# National Institute for Health and Care Excellence

Draft for consultation

# Post-traumatic stress disorder: management (update)

[B] Evidence reviews for psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children

NICE guideline <number>
Evidence reviews

June 2018

**Draft for Consultation** 

These evidence reviews were developed by the National Guideline Alliance hosted by the Royal College of Obstetricians and Gynaecologists



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# Psychological, psychosocial and other

# 2 non-pharmacological interventions for

- 3 the treatment of PTSD in children and
- 4 young people
- 5 This evidence report contains information on 1 review relating to the treatment of PTSD.
- Review question 1.2 For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?

- 1 Review question For children and young people with
- 2 clinically important post-traumatic stress symptoms,
- 3 what are the relative benefits and harms of
- 4 psychological, psychosocial or other non-
- 5 pharmacological interventions targeted at PTSD
- 6 symptoms?
- 7 Summary of the protocol (PICO table)
- 8 Please see Table 1 for a summary of the Population, Intervention, Comparison and
- 9 Outcome (PICO) characteristics of this review.

#### 10 Table 1: Summary of the protocol (PICO table)

| Table 1: Summary of the protocol (PICO table) |   |  |
|---|---|--|
| Population                                    | Children and young people (under 18 years) with clinically important post-traumatic stress symptoms (more than one month after a traumatic event), defined by a diagnosis of PTSD according to DSM, ICD or similar criteria (including PTSD for children 6 years and younger) or clinically-significant PTSD symptoms as indicated by baseline scores above threshold on a validated scale  |  |
| Intervention                                  | Psychological interventions (psychological interventions listed below are examples of interventions which may be included either alone or in combination and delivered to the child or young person and/or a parent or carer in an individual or group format):  • Trauma-focused cognitive behavioural therapies (CBT), including cognitive therapy, cognitive processing therapy, compassion focused therapy, exposure therapy/prolonged exposure (PE), virtual reality exposure therapy (VRET), imagery rehearsal therapy, mindfulness-based cognitive therapy (MBCT) and narrative exposure therapy for traumatized children and adolescents (KidNET)  • Non-trauma-focused CBT, including stress inoculation training (SIT)  • Psychologically-focused debriefing (including single session debriefing)  • Eye movement desensitisation and reprocessing (EMDR)  • Hypnotherapy  • Psychodynamic therapies, including traumatic incident reduction (TIR)  • Counselling, including non-directive/supportive/person-centred counselling  • Human givens therapy  • Combined somatic and cognitive therapies, including thought field therapy (TFT) and emotional freedom technique (EFT)  • Parent training/family interventions, including behavioural family therapy (such as Child and Family Traumatic Stress Intervention [CFTSI]) |  |

|            | <ul> <li>Psychosocial interventions (psychosocial interventions listed below are examples of interventions which may be included either alone or in combination):</li> <li>Meditation</li> <li>Mindfulness-based stress reduction (MBSR)</li> <li>Nature-assisted therapies (including ecotherapy, horticultural therapy, therapeutic horticulture and nature-based therapy)</li> <li>Supported employment (including individual placement and support [IPS] supported employment and Veterans Health Administration Vocational Rehabilitation Programme [VRP])</li> <li>Practical support (including financial and housing)</li> <li>Psychoeducational interventions</li> <li>Peer support (including self-help groups and support groups)</li> </ul> |
|------------|--|
|            | Other non-pharmacological interventions (other non-pharmacological interventions listed below are examples of interventions which may be included either alone or in combination):  • Acupuncture (including classical acupuncture, electroacupuncture, auricular acupuncture, laser acupuncture   |
|            | <ul> <li>and acupoint stimulation [such as acupressure, moxibustion and tapping])</li> <li>Exercise (including anaerobic [such as heavy weight training, sprinting, high-intensity interval training] and aerobic [such as running/jogging, swimming, cycling and walking] exercise, both supervised and unsupervised)</li> <li>Repetitive transcranial magnetic stimulation (rTMS)</li> </ul>   |
| Comparison | <ul> <li>Yoga (including all types of yoga)</li> <li>Any other intervention</li> <li>Treatment as usual</li> <li>Waitlist</li> <li>Placebo</li> </ul>  |
| Outcome    | <ul> <li>Critical outcomes:</li> <li>Efficacy (PTSD symptoms/diagnosis/response/remission /relapse)</li> <li>Acceptability of the intervention (discontinuation for any reason used as a proxy)</li> <li>Important outcomes:</li> <li>Dissociative symptoms</li> <li>Personal/social/educational functioning (including global functioning/functional impairment)</li> <li>Sleeping difficulties</li> <li>Quality of life</li> <li>Symptoms of a coexisting condition (including anxiety, depression and emotional and behavioural problems)</li> </ul>  |

2 For full details see review protocol in Appendix A.

# Psychological interventions for the treatment of PTSD in children and young people

#### 3 Introduction

- 4 A significant proportion of children and young people exposed to potentially traumatic
- 5 events will develop clinically significant symptoms of PTSD, and these symptoms
- 6 may fulfil the diagnostic criteria for PTSD (Alisic 2014). Furthermore, research
- 7 demonstrates that children and young people who have PTSD six months after the
- 8 traumatic event(s) occurred are very unlikely to recover without intervention (Hiller
- 9 2016). This chapter, which informed and steered the recommendations made in the
- 10 updated guideline, reviews research evidence which examines the impact of
- psychological, psychosocial and other non-pharmacological interventions on PTSD
- 12 symptoms.
- 13 Psychological interventions will be considered as classes of intervention (trauma-
- focused CBT; non-trauma-focused CBT; psychologically-focused debriefing; eye
- movement desensitisation and reprocessing [EMDR]; hypnotherapy; psychodynamic
- therapies; counselling; combined somatic and cognitive therapies; parent
- training/family interventions; play therapy; self-help [without support]) and form the
- 18 subsections below.
- 19 Evidence for humans givens therapy was also searched for but none was found.

#### 20 Methods and processes

- 21 This evidence review was developed using the methods and processes as described
- in Developing NICE guidelines: the manual; see the methods chapter for further
- 23 information.
- 24 Declarations of interest were recorded according to NICE's 2014 and 2018 conflicts
- of interest policies.

#### 26 Trauma-focused cognitive behavioural therapies (CBT): clinical evidence

#### 27 Included studies

- 28 Eighty-seven studies of trauma-focused CBT for the treatment of PTSD in children
- and young people were identified for full-text review. Of these 87 studies, 28 RCTs
- 30 (N=2301) were included. Some of these 87 RCTs were three- or four-armed trials
- 31 and as such were included in more than one comparison. There were 8 comparisons
- 32 for trauma-focused CBT.
- 33 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 34 symptoms, there was evidence for 1 relevant comparison: 1 RCT (N=31) compared
- 35 trauma-focused CBT with meditation (Catani 2009/ Rockstroh & Schauer 2004
- 36 [published paper and protocol]).
- For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 38 PTSD symptoms, 19 RCTs (N=1470) compared trauma-focused CBT with waitlist,
- 39 TAU or no treatment (Ahrens & Rexford 2002; Al-Hadethe 2015; Auslander 2017;
- 40 Berger & Gelkopf 2009; Chen 2014; de Roos 2017; Deblinger 1996/ Deblinger 1999
- 41 [one study reported across two papers]; Ertl 2011/Neuner 2007 [published paper and
- 42 protocol]; Goldbeck 2016/ Sachser 2016 [one study reported across two papers];
- Jaycox 2009; Jensen 2014/2017 [one study reported across two papers]; King 2000;

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- 1 Langley 2015; Meiser-Stedman 2010/Meiser-Stedman 2017 [protocol and published
- 2 paper]; Pityaratstian 2015; Ruf 2010; Shein-Szydlo 2016; Smith 2007; Stein
- 3 2003a/Kataoka 2011 [one study reported across two papers]). 8 RCTs (N=718)
- 4 compared trauma-focused CBT with supportive counselling (Chen 2014; Cohen &
- 5 Mannarino 1998/Cohen 2005a [one study reported across two papers]; Cohen
- 6 2004a/Deblinger 2006 [one study reported across two papers]; Cohen 2011/Cohen
- 7 2005b [published paper and protocol]; Ertl 2011/Neuner 2007 [published paper and
- 8 protocol]; Foa 2013a/McLean 2015a/Capaldi 2016/Kaczkurkin 2016/Zandberg. 2016
- 9 [one study reported across five papers]; Ford 2012; Gilboa-Schechtman & Foa
- 10 2004/Gilboa-Schechtman 2010 [protocol and published paper]). 2 RCTs (N=151)
- 11 compared trauma-focused CBT with EMDR (de Roos 2017; Diehle 2015/Lindauer
- 12 2009 [published paper and protocol]). 1 RCT (N=60) compared trauma-focused CBT
- 13 with emotional freedom technique (EFT) (Al-Hadethe 2015). 1 RCT (N=36) compared
- 14 a combined trauma-focused CBT and parent training intervention with waitlist (King
- 15 2000). 1 RCT (N=100) compared trauma-focused CBT with parent training (CBT with
- 16 parent-only) (Deblinger 1996/1999 [one study reported across two papers]). 1 RCT
- 17 (N=159) compared trauma-focused CBT in addition to a psychoeducational group
- 18 with a psychoeducational group-only (Layne 2008).
- 19 Sub-analyses were possible for the trauma-focused CBT versus waitlist, TAU or no
- 20 treatment, and trauma-focused CBT versus supportive counselling comparisons,
- 21 comparing effects by multiplicity of trauma, specific intervention, format, age range,
- 22 diagnostic status at baseline, and trauma type.

#### 23 Excluded studies

- 24 Fifty-nine studies were reviewed at full text and excluded from this review. The most
- 25 common reasons for exclusion were the paper was a systematic review with no new
- 26 useable data and any meta-analysis results not appropriate to extract, or a subgroup
- 27 or secondary analysis of an RCT already included, or the study was unpublished
- 28 (registered on clinical trials.gov and author contacted for full trial report but not
- 29 provided).
- 30 Studies not included in this review with reasons for their exclusions are provided in
- 31 Appendix K.

#### 32 Summary of clinical studies included in the evidence review

- 33 See also the study selection flow chart in Appendix C, forest plots in Appendix E and 34 study evidence tables in Appendix D.
- 35 Table 2, NR - Not reported
- 36 Table 3, ADHD, Attention Deficit Hyperactivity Disorder; NR, not reported; TAU, Treatment as usual;
- 37 TF-CBT, trauma-focused CBT; PTSD, Post-traumatic stress disorder; OCD, Obsessive Compulsive
- 38 Disorder; SSET, Support for Students Exposed to Trauma; CT-PTSD, Cognitive Therapy Post-traumatic
- 39 stress disorder; TRT, Teaching Recovery Techniques; CBT, Cognitive Behavioural Therapy.
- 40 <sup>1</sup>Ahrens 2002; <sup>2</sup>Al-Hadethe 2015; <sup>3</sup>Auslander 2017; <sup>4</sup>Berger 2009; <sup>5</sup>Chen 2014; <sup>6</sup>de Roos 2017;
- <sup>7</sup>Deblinger 1996/1999; <sup>8</sup>Ertl 2011/Neuner 2007; <sup>9</sup>Goldbeck 2016/Sachser 2016; <sup>10</sup>Jaycox 2009; <sup>11</sup>Jensen 2014/2017; <sup>12</sup>King 2000; <sup>13</sup>Langley 2015; <sup>14</sup>Meiser-Stedman 2010/2017; <sup>15</sup>Pityaratstian 2015; <sup>16</sup>Ruf 41
- 42
- 43 2010; <sup>17</sup>Shein-Szydlo 2016; <sup>18</sup>Smith 2007; <sup>19</sup>Stein 2003a/Kataoka 2011

44

- 45 Table 4 and CBT, Cognitive Behavioural Therapy; EMDR, Eye Movement Desensitisation and
- 46 Reprocessing; EFT, Emotional Freedom Techniques; PTSD – Post-traumatic stress disorder; NR, Not
- 47 relevant; ICD/ DSM, International Classification of Disease/ Diagnostic and Statistical Manual of Mental
- 48 Disorders; SAS-CBT, Sexual abuse specific cognitive behavioural therapy.
- 49 <sup>1</sup>Chen 2014; <sup>2</sup>Cohen 1998/2005a; <sup>3</sup>Cohen 2004a/Deblinger 2006; <sup>4</sup>Cohen 2011/2005b; <sup>5</sup>Ertl

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- 2011/Neuner 2007; <sup>6</sup>Foa 2013a/McLean 2015a/Capaldi 2016/Kaczkurkin 2016/Zandberg 2016; <sup>7</sup>Ford
   2012; <sup>8</sup>Gilboa-Schechtman 2004/2010; <sup>9</sup>de Roos 2017; <sup>10</sup>Diehle 2015/Lindauer 2009
- Table 5 provide brief summaries of the included studies and evidence from these are
- 4 summarised in the clinical GRADE evidence profiles below (Table 6, Table 7, Table
- 5 8, Table 9, Table 10, Table 11, Table 12 and Table 13).
- See also the study selection flow chart in Appendix C, forest plots in Appendix E and study evidence tables in Appendix D.

## Table 2: Summary of included studies: Trauma-focused CBT for early treatment (1-3 months)

| Comparison                               | Trauma-focused CBT versus meditation   |
|--|--|
|  | Tradina rocacca CDT refeat incaration  |
| Total no. of studies (N randomised)      | 1 (31)   |
| Study ID                                 | Catani 2009/Rockstroh 2004   |
| Country                                  | Sri Lanka  |
| Diagnostic status                        | Clinically important PTSD symptoms (scoring above a threshold on validated scale)  |
| Mean months since onset of PTSD          | NR   |
| Mean age (range)                         | 11.9 (8-14)  |
| Sex (% female)                           | 45   |
| Ethnicity (% BME)                        | NR   |
| Coexisting conditions                    | NR   |
| Mean months since traumatic event        | Mean NR (study carried out within the first months after the tsunami disaster in Sri Lanka)  |
| Type of traumatic event                  | Natural disaster: Tsunami disaster in Sri Lanka  |
| Single or multiple incident index trauma | Single   |
| Lifetime experience of trauma            | Mean number of traumas 4.6. 81% identified the tsunami as the worst traumatic event experienced but 68% had also been affected by traumatic war experiences  |
| Intervention details                     | Narrative exposure therapy for traumatized children and adolescents (KidNET)   |
| Intervention format                      | Individual   |
| Intervention intensity                   | 6x thrice-weekly 60-90-min sessions (6-9 hours)  |
| Comparator                               | Meditation-relaxation, sessions containing meditation and relaxation techniques including 'inner peace meditation', 'uchchadana mantra chanting', 'progressive muscle relaxation', 'ice cream body relaxation', and 'inner light meditation' |
| Intervention length (weeks)              | 2  |

10 NR – Not reported

11

12

8

9

# Table 3: Summary of included studies: Trauma-focused CBT for delayed treatment (>3 months)-part 1

| tication (* 6 months)-part i        |   |
|-------------------------------------|---|
| Comparison                          | Trauma-focused CBT versus waitlist, TAU or no treatment |
| Total no. of studies (N randomised) | 19 (1470)   |
| Study ID                            | Ahrens 2002 <sup>1</sup>                                |

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|                                 | Trauma focused CPT versus weither TALL or no  |
|---------------------------------|---|
| Comparison                      | Trauma-focused CBT versus waitlist, TAU or no treatment   |
|                                 | Al-Hadethe 2015 <sup>2</sup> Auslander 2017 <sup>3</sup> Berger 2009 <sup>4</sup> Chen 2014 <sup>5</sup> de Roos 2017 <sup>6</sup> Deblinger 1996/1999 <sup>7</sup> Ertl 2011/Neuner 2007 <sup>8</sup> Goldbeck 2016/Sachser 2016 <sup>9</sup> Jaycox 2009 <sup>10</sup> Jensen 2014/2017 <sup>11</sup> King 2000 <sup>12</sup> Langley 2015 <sup>13</sup> Meiser-Stedman 2010/2017 <sup>14</sup> Pityaratstian 2015 <sup>15</sup> Ruf 2010 <sup>16</sup> Shein-Szydlo 2016 <sup>17</sup> Smith 2007 <sup>18</sup> Stein 2003a/Kataoka 2011 <sup>19</sup> |
| Country                         | US1,3,7,10,13,19 Iraq <sup>2</sup> Sri Lanka <sup>4</sup> China <sup>5</sup> Netherlands <sup>6</sup> Uganda <sup>8</sup> Germany <sup>9,16</sup> Norway <sup>11</sup> Australia <sup>12</sup> UK <sup>14,18</sup> Thailand <sup>15</sup> Mexico <sup>17</sup>  |
| Diagnostic status               | PTSD diagnosis according to ICD/DSM criteria <sup>1,2,8,14,15,16,17</sup> Clinically important PTSD symptoms (scoring above a threshold on validated scale) <sup>3,4,5,6,7,9,10,11,12,13,18,19</sup>  |
| Mean months since onset of PTSD | NR  |
| Mean age (range)                | 16.4 (15-18) <sup>1</sup> Mean NR (16-19) <sup>2</sup> 14.6 (12-18) <sup>3</sup> Mean NR (9-14) <sup>4</sup> 14.5 (range NR) <sup>5</sup> 13.1 (8-18) <sup>6</sup> 9.8 (7-13) <sup>7</sup> 18.4 (12-25) <sup>8</sup> 13 (7-17) <sup>9</sup> 11.5 (range NR) <sup>10</sup> 15.1 (10-18) <sup>11</sup> 11.4 (5-17) <sup>12</sup> 7.7 (6-11) <sup>13</sup>   |

|                       | Trauma-focused CBT versus waitlist, TAU or no   |
|-----------------------|---|
| Comparison            | treatment   |
|                       | 13.3 (8-17) <sup>14</sup> 12.3 (10-15) <sup>15</sup> 11.4 (7-16) <sup>16</sup> 14.9 (12-18) <sup>17</sup> 13.9 (range NR) <sup>18</sup> 11 (range NR) <sup>19</sup>   |
| Sex (% female)        | 01.2<br>1003<br>484<br>685<br>576<br>837<br>558<br>729<br>5110<br>8011<br>6912<br>5013,18<br>7214,15<br>4616<br>6417<br>5619  |
| Ethnicity (% BME)     | 40 <sup>1</sup> NR <sup>2,4,5,6,8,9,11,12,15,16,17,19</sup> 78 <sup>3</sup> 28 <sup>7</sup> 96 <sup>10</sup> 73 <sup>13</sup> 14 <sup>14</sup> 54 <sup>18</sup>   |
| Coexisting conditions | 52% stated they had experienced a head injury that led to loss of consciousness; 40% stated that they had been diagnosed with ADD or ADHD in the past¹ NR²,3,4,5,8,10,11,13,15,16,19 54% had one or more co-morbid disorder (assessed with ADIS-C) 6 29% major depression; 30% oppositional defiant disorder; 20% ADHD; 11% separation anxiety; 6% conduct disorder; 5% specific phobia; 1% OCD <sup>7</sup> 34% >1 comorbid DSM-IV disorder: Depressive disorders (20%); Anxiety disorders (10%); ADHD (6%); Disruptive behaviour disorders (4%)9 For 69% who met DSM-IV criteria for full PTSD (N=25): 16% with full PTSD had no other Axis I diagnoses, 36% had one comorbid diagnosis, 40% had two comorbid diagnoses, and 8% had three comorbid diagnoses. The comorbid diagnoses included dysthymia (28%), oppositional defiant disorder (28%), separation anxiety disorder (24%), generalized anxiety disorder (20%), conduct disorder (12%), major depression (8%), attention-deficit/hyperactivity disorder (8%), and specific phobia (8%)¹² |

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|                                   | Transport for a second ODT reserves are idlied. TALL on the  |
|-----------------------------------|--|
| Comparison                        | Trauma-focused CBT versus waitlist, TAU or no treatment  |
|                                   | 86% comorbid anxiety disorder; 55% comorbid affective disorder; 52% comorbid behavioural disorder <sup>14</sup> 14% anxiety disorder; 28% depression <sup>17</sup> 79% had any comorbidity <sup>18</sup>   |
| Mean months since traumatic event | NR <sup>1,2,3,10,11,13,17,19</sup> NR (around 24 months) <sup>4</sup> 24 <sup>5</sup> 16.5 <sup>6</sup> Mean NR (for 66% the last sexually abusive incident occurred in the 6 months prior to initial assessment, 16% 6 months to 2 years before initial assessment, and 18% 2 more years before the evaluation) <sup>7</sup> 80.5 <sup>8</sup> NR (inclusion criteria >3 months) <sup>9</sup> 54.5 <sup>12</sup> 3.9 <sup>14</sup> >48 <sup>15</sup> Mean NR (37.25 months in exile) <sup>16</sup> Mean NR (median: 8.65; range: 3.3-64 months) <sup>18</sup>   |
| Type of traumatic event           | Mixed: Adolescent offenders incarcerated in a youth facility. 68% had documented trauma histories (as documented in their charts from collateral sources ranging from Social Rehabilitation Service investigations, child protective services reports, hospital reports, etc.) 1 Unclear (no details reported) 2 Mixed: Girls involved in child welfare who had histories of abuse and neglect. Girls with histories of sexual abuse were included3 Natural disaster: Tsunami (Sri Lanka, December 26 2004) -84% present and physically hurt during the tsunami; 12% present during the tsunami, but were not hurt; 4% not personally exposed to the tsunami4 Natural disaster: Adolescents who had lost at least one parent in the Sichuan, China, Earthquake5 Mixed: Physical abuse/assault (23%); Sexual abuse (26%); Accident/injury of a loved one (19%); Traumatic loss (18%); Disaster/other (13%)6 Childhood sexual abuse: Contact sexual abuse. 18% experienced 1 sexually abusive incident, 47% 2-10 episodes, 22% 11-50 episodes, and 13% >50 abusive incidents7 Child soldiers: The duration of abduction ranged from several hours to 7.42 years, with a median of 2.47 months. The likelihood of an event being indicated as the worst if present was highest for being forced to kill (55%), followed by witnessed killing (31%) and seeing someone being mutilated or seeing dead bodies (13%)8 Mixed: Interpersonal trauma (77%); accidental (23%). The most frequently reported traumatic index events were experiences of sexual abuse, sexual assaults, physical violence, or witnessing domestic violence9 Exposure to non-sexual violence: Experience of severe violence in the prior year <sup>10</sup> |

|  | Trauma-focused CBT versus waitlist, TAU or no   |
|--|---|
| Comparison                               | treatment   |
| Comparison                               | · · · · · · · · · · · · · · · · · · ·   |
|  | earthquakes (19%), other natural disasters (12%), and sexual abuse (8%) <sup>16</sup> Mixed: Street Children in Mexico City - 56% were victims of sexual abuse,47% of physical abuse, 18% of witnessing a violent event, and 17% of death of a family member <sup>17</sup> Motor Vehicle Collision: Motor vehicle accident (50%); |
|  | Assault (38%); Witnessed violence (13%) <sup>18</sup> Exposure to non-sexual violence: 76% any violence involving a gun or knife. Number of violent events experienced: 2.8; Number of violent events witnessed: 5.95 <sup>19</sup>   |
| Single or multiple incident index trauma | Multiple <sup>1,3,7,8,9,11,12,13,16,17,19</sup> Unclear <sup>2</sup> Single <sup>4,5,6,10,14,15,18</sup>  |
| Lifetime experience of trauma            | 29% had experienced multiple traumas <sup>1</sup> NR <sup>2,3,5,6,7,10,13,15,19</sup>   |

|                      | Trauma focused CRT versus waitlist TALL or no   |
|----------------------|---|
| Comparison           | Trauma-focused CBT versus waitlist, TAU or no treatment   |
|                      | 89% had been exposed to a major traumatic incident not related to the tsunami <sup>4</sup>  |
|                      | Other than abduction, the most common traumatic event types reported by 81 or more of the 85 participants were exposure to a war zone, witnessing someone being killed, witnessing abduction, witnessing physical assault, and assaults with weapons <sup>8</sup> |
|                      | Number of traumatic events: 6.35 (SD=3.70) 9  |
|                      | Mean 3.6 different types of traumas (SD=1.8, range=1–10) 11   |
|                      | Mean number of abusive episodes: 7.6 (SD=3.8; range 1-33) <sup>12</sup>   |
|                      | 38% had experienced previous trauma <sup>14</sup>   |
|                      | Mean number of traumatic event types: 4.4 <sup>16</sup>   |
|                      | 35% reported more than one type of traumatic event <sup>17</sup> 29% prior exposure to trauma <sup>18</sup>   |
| Intervention details | Cohen TF-CBT/Cognitive processing therapy: Cognitive processing therapy (following the manual by Resick & Schnicke 1993) <sup>1</sup>   |
|                      | Narrative exposure therapy for traumatized children and adolescents (KidNET) following protocol of Neuner (2008) <sup>2,16</sup>  |
|                      | GAIN (Girls Aspiring Toward Independence); an adapted form of CBITS (Cognitive Behavioural Intervention for Trauma in Schools), developed with input from focus groups, caregivers and other involved stakeholders <sup>3</sup>                                   |
|                      | ERASE Stress Sri Lanka (ES-SL; following manual of Berger & Manasra, 2005) 4  |
|                      | Adapted Teaching Recovery Techniques group CBT intervention (Smith, Dyregrov, Yule 1999) <sup>5</sup>   |
|                      | Cognitive behavioural writing therapy (CBWT; following manual by Van der Oord 2010) <sup>6</sup>  |
|                      | Exposure therapy (following manual by Deblinger & Heflin, 1996) <sup>7,12</sup>   |
|                      | Narrative Exposure Therapy (kidNET) adapted for the field8  |
|                      | Cohen TF-CBT/Cognitive processing therapy (according to Cohen's 2006 manual) with parallel or conjoint sessions with child and caregiver <sup>9,11</sup>  |
|                      | Support for Students Exposed to Trauma (SSET) adapted from the Cognitive Behavioural Intervention for Trauma in Schools (CBITS) programme <sup>10</sup>   |
|                      | Bounce Back, Trauma-focused CBT intervention in school setting and involving caregivers <sup>13</sup>   |
|                      | Cognitive Therapy for PTSD (CT-PTSD), based upon the treatment manual from Smith (2010) <sup>14</sup>   |
|                      | Brief group CBT, Teaching Recovery Techniques (TRT; Smith 1999), adapted to span 3 days and sessions made longer to accommodate content <sup>15</sup>   |
|                      | Cohen TF-CBT/Cognitive processing therapy: CBT for Trauma in Street Children <sup>17</sup>  |
|                      | Cognitive therapy based on protocol from Ehlers (2000) with adaptations (Yule 2005) for children <sup>18</sup>  |
|                      | Cognitive behavioural intervention for trauma in schools (CBITS; following manual of Jaycox 2003) 19  |

|                                   | Trauma-focused CBT versus waitlist, TAU or no   |
|-----------------------------------|---|
| Comparison                        | treatment   |
| Intervention format               | Individual <sup>1,2,6,7,8,12,14,16,17</sup>   |
|                                   | Group <sup>3,4,5,10,13,15,19</sup>  |
|                                   | Individual/Family <sup>9,11,18</sup>  |
| Intervention intensity            | 8x 1-hour sessions (8 hours) 1  |
|                                   | 4x biweekly 60-90 min sessions (4-6 hours) <sup>2</sup>   |
|                                   | 10x 90 min sessions (plus pre- and post-intervention  |
|                                   | 'parties'), plus 2x supportive adult sessions <sup>3</sup>  |
|                                   | 12x weekly 90-min sessions (18 hours in total) 4  |
|                                   | 6x weekly 1-hour sessions (6 hours) <sup>5</sup> 6x weekly 45-min sessions (4.5 hours). Mean attended 5.4                       |
|                                   | (SD=0.78) sessions <sup>6</sup>   |
|                                   | 12x weekly 45-min sessions (9 hours) <sup>7</sup>   |
|                                   | 8x thrice-weekly 90-120-min sessions (12-16 hours) <sup>8</sup>   |
|                                   | 12x weekly 90-min sessions (18 hours). 86% completed at   |
|                                   | least 8 sessions and 76% completed the full 12 sessions <sup>9</sup>  |
|                                   | 10x 45-min sessions (7.5 hours) 10  |
|                                   | 12-15x sessions (length of session NR) 11   |
|                                   | 20x weekly 50-min sessions (16.7 hours) 12  |
|                                   | 10x 50-60min child group sessions, 2-3x 30-50min child sessions and 1-3x 30-50min caregiver sessions <sup>13</sup>              |
|                                   | 10x weekly 90-min sessions (15 hours). Mean attended 8.3 sessions (SD=2.2) 14   |
|                                   | 3x 2-hour sessions (6 hours) 15   |
|                                   | 8x weekly sessions (length of sessions NR) 16   |
|                                   | 12x weekly 1-hour sessions (12 hours) 17  |
|                                   | 10x weekly sessions (length of session NR) 18,19  |
| Comparator                        | Waitlist <sup>1,4,6,8,9,10,12,13,14,15,16,17,18,19</sup>  |
|                                   | No treatment <sup>2,5</sup>   |
|                                   | TAU: Any care required other than the experimental intervention <sup>3</sup>  |
|                                   | TAU: Community control, parents and children were given   |
|                                   | information about symptom patterns and encouraged to  |
|                                   | access therapy, and child protection workers or the victim witness coordinator were asked to assist with referrals <sup>7</sup> |
|                                   | TAU: Clinician asked to provide the treatment they felt would   |
|                                   | be effective. All participants received individual treatment (no  |
|                                   | group treatment), but in 55% of the cases, parents were also  |
|                                   | involved in the therapy process <sup>11</sup>   |
| Intervention length (weeks)       | 81,16   |
|                                   | 22  |
|                                   | 133,13  |
|                                   | 124,7,9,17  |
|                                   | 6 <sup>5,6</sup><br>3 <sup>8</sup>  |
|                                   | 1010,14,18,19   |
|                                   | NR <sup>11</sup>  |
|                                   | 20 <sup>12</sup>  |
|                                   | 0.4 <sup>15</sup>   |
| ADID Attention Definit II may not | ivity Disorder: NR_not renorted: TALL Treatment as usual: TF-CRT  |

ADHD, Attention Deficit Hyperactivity Disorder; NR, not reported; TAU, Treatment as usual; TF-CBT, trauma-focused CBT; PTSD, Post-traumatic stress disorder; OCD, Obsessive Compulsive Disorder;

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SSET, Support for Students Exposed to Trauma; CT-PTSD, Cognitive Therapy Post-traumatic stress disorder; TRT, Teaching Recovery Techniques; CBT, Cognitive Behavioural Therapy.

¹Ahrens 2002; ²Al-Hadethe 2015; ³Auslander 2017; ⁴Berger 2009; ⁵Chen 2014; ⁶de Roos 2017; ¹Deblinger 1996/1999; ⁶Ertl 2011/Neuner 2007; ⁶Goldbeck 2016/Sachser 2016; ¹⁰Jaycox 2009; ¹¹Jensen 2014/2017; ¹²King 2000; ¹³Langley 2015; ¹⁴Meiser-Stedman 2010/2017; ¹⁵Pityaratstian 2015; ¹⁶Ruf 2010; ¹³Shein-Szydlo 2016; ¹⁶Smith 2007; ¹९Stein 2003a/Kataoka 2011

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# Table 4: Summary of included studies: Trauma-focused CBT for delayed treatment (>3 months)-part 2

| treatment (>3 months)-part 2             |   |   |  |
|--|---|---|--|
| Comparison                               | Trauma-focused CBT versus supportive counselling  | Trauma-focused CBT versus EMDR  | Trauma-focused CBT versus EFT                |
| Total no. of studies (N randomised)      | 8 (718)   | 2 (151)   | 1 (60)                                       |
| Study ID                                 | Chen 2014 <sup>1</sup> Cohen 1998/2005a <sup>2</sup> Cohen 2004a/ Deblinger 2006 <sup>3</sup> Cohen 2011/2005b <sup>4</sup> Ertl 2011/Neuner 2007 <sup>5</sup> Foa 2013a/McLean 2015a/Capaldi 2016/Kaczkurkin 2016/Zandberg 2016 <sup>6</sup> Ford 2012 <sup>7</sup> Gilboa-Schechtman 2004/2010 <sup>8</sup> | de Roos 2017 <sup>9</sup> Diehle 2015/Lindauer 2009 <sup>10</sup>                             | Al-Hadethe 2015                              |
| Country                                  | China <sup>1</sup><br>US <sup>2,3,4,6,7</sup><br>Uganda <sup>5</sup><br>Israel <sup>8</sup>   | Netherlands   | Iraq   |
| Diagnostic<br>status                     | Clinically important PTSD symptoms (scoring above a threshold on validated scale) 1,2,4,7 PTSD diagnosis according to ICD/DSM criteria <sup>3,5,6,8</sup>   | Clinically important<br>PTSD symptoms<br>(scoring above a<br>threshold on validated<br>scale) | PTSD diagnosis according to ICD/DSM criteria |
| Mean<br>months<br>since onset<br>of PTSD | NR  | NR  | NR   |
| Mean age<br>(range)                      | 14.5 (range NR) <sup>1</sup> 11.1 (7-15) <sup>2</sup> 10.8 (8-14) <sup>3</sup> 9.6 (7-14) <sup>4</sup> 18.4 (12-25) <sup>5</sup> 15.3 (13-18) <sup>6</sup> 14.7 (13-17) <sup>7</sup> 14.1 (12-18) <sup>8</sup>  | 13.1 (8-18) <sup>9</sup><br>12.9 (8-18) <sup>10</sup>   | Mean NR (16-19)                              |

| Comparison                                    | Trauma-focused CBT versus supportive counselling  | Trauma-focused CBT versus EMDR   | Trauma-focused CBT versus EFT |
|---|---|--|-------------------------------|
| Sex (% female)                                | 68 <sup>1</sup> 69 <sup>2</sup> 79 <sup>3</sup> 51 <sup>4</sup> 55 <sup>5</sup> 100 <sup>6,7</sup> 63 <sup>8</sup>  | 57 <sup>9</sup><br>62 <sup>10</sup>  | 0                             |
| Ethnicity (%<br>BME)                          | NR <sup>1,5,8</sup> 41 <sup>2</sup> 40 <sup>3</sup> 44 <sup>4</sup> 82 <sup>6</sup> 75 <sup>7</sup>   | NR   | NR                            |
| Coexisting conditions                         | NR <sup>1,2,3,4,5</sup> 57% had ≥1 comorbid psychiatric diagnoses <sup>6</sup> 34% major depressive disorder, 26% oppositional defiant disorder, 23% conduct disorder, and 13% attention deficit hyperactivity disorder <sup>7</sup> 81% ≥ 1 comorbid disorder: 50% had one additional internalizing disorder, 13% had an additional externalizing disorder, and 16% had internalizing and externalizing disorders <sup>8</sup> | 54% had one or more<br>co-morbid disorder<br>(assessed with ADIS-<br>C) <sup>9</sup><br>NR <sup>10</sup>   | NR                            |
| Mean<br>months<br>since<br>traumatic<br>event | Mean NR (inclusion criteria within 6 months) <sup>2</sup> 12.3 <sup>3</sup> NR (IPV duration: 5% <2 years, 19% 2-5 years, 77% >5 years) <sup>4</sup> 80.5 <sup>5</sup> 40.5 <sup>6</sup> NR <sup>7</sup> 18.5 <sup>8</sup>  | 16.5 <sup>9</sup><br>NR <sup>10</sup>  | NR                            |
| Type of traumatic event                       | Natural disaster: Adolescents who had lost at least one parent in the Sichuan, China, Earthquake <sup>1</sup> Childhood sexual abuse: Contact sexual abuse perpetrated by   | Mixed: Physical<br>abuse/assault (23%);<br>Sexual abuse (26%);<br>Accident/injury of a<br>loved one (19%);<br>Traumatic loss (18%);<br>Disaster/other (13%) <sup>9</sup> | Unclear (details NR)          |

|            | Trauma francis I ODT   | Trauma forma I ODT   | Trauma former LODE            |
|------------|--|--|-------------------------------|
|            | Trauma-focused CBT versus supportive   | Trauma-focused CBT versus EMDR   | Trauma-focused CBT versus EFT |
| Comparison | counselling  |  |                               |
|            | someone at least 5 years older than the participants (36% single episode, 21% 2- 5 abuse occasions, 8% 6-10 times, 33% were abused more than 10 times; 2% unknown) 2 Childhood sexual abuse: Contact sexual abuse: Contact sexual abuse <sup>3</sup> Domestic violence: Children exposed to intimate partner violence <sup>4</sup> Child soldiers: The duration of abduction ranged from several hours to 7.42 years, with a median of 2.47 months. The likelihood of an event being indicated as the worst if present was highest for being forced to kill (55%), followed by witnessed killing (31%) and seeing someone being mutilated or seeing dead bodies (13%) <sup>5</sup> Childhood sexual abuse <sup>6</sup> Mixed: Trauma exposure was extensive, including 97% to a traumatic accident, disaster, or illness; 88% to physical assault or abuse; 81% to traumatic community violence; 78% to traumatic family violence; 78% to traumatic family violence; 44% to sexual assault or abuse; 41% to traumatic emotional abuse; and 29% to traumatic bullying <sup>7</sup> Mixed: Terrorist attack (13%); motor vehicle accident (42%); non- sexual assault (21%); Other (18%) <sup>8</sup> | Mixed: 63% Single-event index trauma. Single event traumas: accidents (23 %), sexual assault (17 %); threat (with weapon) (13 %); kidnapping (10 %); serious illness (7 %); or other (30 %). Multiple-event traumas: exposure to domestic violence (44 %) and sexual assault (39 %) and other (17 %) <sup>10</sup> |                               |

| Comparison  | Trauma-focused CBT versus supportive counselling  | Trauma-focused CBT versus EMDR  | Trauma-focused CBT versus EFT  |
|---|---|---|--|
| Single or<br>multiple<br>incident<br>index trauma | Single <sup>1,8</sup> Multiple <sup>2,3,4,5,6,7</sup>   | Single  | Unclear  |
| Lifetime<br>experience<br>of trauma               | NR <sup>1,2,6,7,8</sup> Mean 2.66 (SD 1.61) traumatic events in addition to sexual abuse <sup>3</sup> Mean number of trauma types: 3.7 <sup>4</sup> Other than abduction, the most common traumatic event types reported by 81 or more of the 85 participants were exposure to a war zone, witnessing someone being killed, witnessing abduction, witnessing physical assault, and assaults with weapons <sup>5</sup>   | NR <sup>9</sup> Mean types of prior trauma 6.5 <sup>10</sup>  | NR   |
| Intervention details                              | Adapted Teaching Recovery Techniques group CBT intervention (Smith, Dyregrov, Yule 1999)   Cohen TF- CBT/Cognitive processing therapy: Sexual Abuse-Specific Cognitive-Behavioural Therapy (SAS-CBT; Cohen 2000)   Narrative exposure therapy (based on protocol from Deblinger & Heflin 1996)   Cohen TF- CBT/Cognitive processing therapy   Narrative Exposure Therapy (kidNET) adapted for the field <sup>5</sup> Prolonged exposure—A program (Foa 2008)   Cohen TF- CBT/Cognitive processing therapy: TARGET intervention (Trauma Affect Regulation: Guide for Education and | Cognitive behavioural writing therapy (CBWT; following manual by Van der Oord 2010) 9 Cohen TF-CBT/Cognitive processing therapy (following protocol by Cohen 2008) 10 | Narrative exposure therapy for traumatized children and adolescents (KidNET) following protocol of Neuner (2008) |

|                        | Trauma-focused CBT  | Trauma-focused CBT   | Trauma-focused CBT                         |
|------------------------|---|--|--|
| Commonicon             | versus supportive   | versus EMDR  | versus EFT                                 |
| Comparison             | Counselling Therapy; Ford & Russo, 2006) <sup>7</sup> Adapted prolonged exposure therapy for adolescents (PE-A) <sup>8</sup>  |  |  |
| Intervention format    | Group <sup>1</sup> Individual/Family <sup>2,3,4</sup> Individual <sup>5,6,7,8</sup>   | Individual <sup>9</sup><br>Individual/Family <sup>10</sup>   | Individual                                 |
| Intervention intensity | 6x weekly 1-hour sessions (6 hours) <sup>1</sup> 12x sessions of 90 mins (45mins child, 45mins carer; 18 hours in total) <sup>2</sup> 12x 90-min sessions (9x 45-min for parent and 45-min for child and 3x 30-min joint parent-child session + 30-min for parent and 30-min for child; total 18 hours). Mean attended sessions 10.5 (SD=2.9) <sup>3</sup> 8x 90-min sessions (45-min for child and 45-min for child and 45-min for parent; 12 hours in total) <sup>4</sup> 8x thrice-weekly 90-120-min sessions (12-16 hours) <sup>5</sup> 14x weekly 60-90 min sessions (14-21 hours). Mean 12 treatment sessions (14-21 hours). Mean 12 treatment sessions 90% attended at least 8 sessions <sup>6</sup> 12x 50-min sessions (10 hours). Mean attended sessions 7 (SD=4.2). 67% at least 5 sessions <sup>7</sup> 12-15x weekly 60-90min sessions (12-22.5 hours). Mean number of sessions 13 and mean therapist hours per patient were 16.88 | 6x weekly 45-min sessions (4.5 hours). Mean attended 5.4 (SD=0.78) sessions <sup>9</sup> 8x weekly 1-hour sessions (8 hours) <sup>10</sup> | 4x biweekly 60-90 min sessions (4-6 hours) |
| Comparator             | General support provided on an individual basis adopting counselling techniques such as   | Eye movement<br>desensitisation and<br>reprocessing (EMDR)<br>based on the standard<br>protocol from Shapiro                               | Emotional Freedom<br>Technique (EFT)       |

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| Comparison                        | Trauma-focused CBT versus supportive counselling   | Trauma-focused CBT versus EMDR   | Trauma-focused CBT versus EFT |
|-----------------------------------|--|--|-------------------------------|
|                                   | listening, reflection, and empathy <sup>1</sup> Nondirective supportive therapy (NST) <sup>2</sup> Client Centred Therapy (CCT; based on unpublished treatment manual) <sup>3,6</sup> Child-centred therapy <sup>4</sup> Needs-based intervention <sup>5</sup> Manualized relational therapy <sup>7</sup> Time-limited Dynamic Psychotherapy for Adolescents (TLDP-A) <sup>8</sup> | (2001) with age-<br>appropriate<br>modifications<br>suggested by Tinker<br>and Wilson (1999) and<br>Greenwald (1999) <sup>9</sup><br>Eye movement<br>desensitisation and<br>reprocessing (EMDR;<br>following Dutch<br>protocol for EMDR for<br>children and<br>adolescents; Beer & de<br>Roos, 2008) <sup>10</sup> |                               |
| Intervention<br>length<br>(weeks) | 6 <sup>1</sup><br>12 <sup>2,3,7</sup><br>8 <sup>4</sup><br>3 <sup>5</sup><br>14 <sup>6</sup><br>15 <sup>8</sup>  | 6 <sup>9</sup><br>8 <sup>10</sup>  | 2                             |

CBT, Cognitive Behavioural Therapy; EMDR, Eye Movement Desensitisation and Reprocessing; EFT, Emotional Freedom Techniques; PTSD – Post-traumatic stress disorder; NR, Not relevant; ICD/ DSM, International Classification of Disease/ Diagnostic and Statistical Manual of Mental Disorders; SAS-CBT, Sexual abuse specific cognitive behavioural therapy.

Sexual abuse specific cognitive behavioural therapy.

¹Chen 2014; ²Cohen 1998/2005a; ³Cohen 2004a/Deblinger 2006; ⁴Cohen 2011/2005b; ⁵Ertl 2011/Neuner 2007; °Foa 2013a/McLean 2015a/Capaldi 2016/Kaczkurkin 2016/Zandberg 2016; ¬Ford 2012; ³Gilboa-Schechtman 2004/2010; ³de Roos 2017; ¹¹Diehle 2015/Lindauer 2009

Table 5: Summary of included studies: Trauma-focused CBT for delayed treatment (>3 months)-part 3

| Comparison                               | Trauma-focused CBT + parent training versus waitlist                              | Trauma-focused CBT versus parent training (CBT with parent-only)                  | Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group     |
|--|---|---|---|
| Total no. of studies (N randomised)      | 1 (36)  | 1 (100)   | 1 (159)   |
| Study ID                                 | King 2000   | Deblinger 1996/1999   | Layne 2008  |
| Country                                  | Australia   | US  | Bosnia  |
| Diagnostic status                        | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Clinically important PTSD symptoms (scoring above a threshold on validated scale) |
| Mean<br>months<br>since onset<br>of PTSD | NR  | NR  | NR  |

|   | Trauma-focused CBT + parent training   | Trauma-focused CBT versus parent training (CBT with parent-only)   | Trauma-focused CBT (+ psychoeducational group) versus psychoeducational   |
|---|--|--|---|
| Comparison                                    | versus waitlist  | 7,   | group   |
| Mean age (range)                              | 11.4 (5-17)  | 9.8 (7-13)   | 16 (13-19)  |
| Sex (% female)                                | 69   | 83   | 64  |
| Ethnicity (%<br>BME)                          | NR   | 28   | NR  |
| Coexisting conditions                         | For 69% who met DSM-IV criteria for full PTSD (N=25): 16% with full PTSD had no other Axis I diagnoses, 36% had one comorbid diagnosis, 40% had two comorbid diagnoses, and 8% had three comorbid diagnoses. The comorbid diagnoses included dysthymia (28%), oppositional defiant disorder (28%), separation anxiety disorder (24%), generalized anxiety disorder (20%), conduct disorder (12%), major depression (8%), attention-deficit/hyperactivity disorder (8%), and specific phobia (8%) | 29% major depression;<br>30% oppositional<br>defiant disorder; 20%<br>ADHD; 11% separation<br>anxiety; 6% conduct<br>disorder; 5% specific<br>phobia; 1% OCD   | NR  |
| Mean<br>months<br>since<br>traumatic<br>event | 54.5   | Mean NR (for 66% the last sexually abusive incident occurred in the 6 months prior to initial assessment, 16% 6 months to 2 years before initial assessment, and 18% 2 more years before the evaluation) | NR  |
| Type of traumatic event                       | Childhood sexual abuse: In the majority of cases, the offenders were male adults known to the child such as the biological father, stepfather, family friend, neighbour, or teacher. Nearly all of the children had  | Childhood sexual abuse: Contact sexual abuse. 18% experienced 1 sexually abusive incident, 47% 2-10 episodes, 22% 11-50 episodes, and 13% >50 abusive incidents  | Witnessing war as a civilian: Approximately 73% of the students participating reported experiencing direct life threat arising from close proximity to exploding shells or rifle fire, 36% reported witnessing during the |

|   |   | Trauma-focused CBT   | Trauma-focused CBT   |
|---|---|--|--|
|   | Trauma-focused CBT  | versus parent<br>training (CBT with                                      | (+ psychoeducational group) versus   |
| Comparison  | + parent training versus waitlist   | parent-only)   | psychoeducational group  |
|   | experienced multiple<br>episodes of sexual<br>abuse involving<br>penetration offenses<br>and other forms of<br>sexual abuse |  | war violent death or serious injury, 12% reported witnessing torture, and 46% reported the serious injury of a person to whom they were close, 14% reported the violent death during the war of a nuclear family member, and 73% reported the violent death of at least one person to whom they were close |
| Single or<br>multiple<br>incident<br>index trauma | Multiple  | Multiple   | Multiple   |
| Lifetime experience of trauma                     | Mean number of abusive episodes: 7.6 (SD=3.8; range 1-33)   | NR   | NR   |
| Intervention details                              | Exposure therapy + parent training  | Exposure therapy<br>(following manual by<br>Deblinger & Heflin,<br>1996) | Trauma and Grief<br>Component Therapy<br>for Adolescents<br>(TGCT)   |
| Intervention format                               | Individual/Family   | Individual   | Group  |
| Intervention intensity                            | 20x weekly 50-min sessions (16.7 hours)   | 12x weekly 45-min sessions (9 hours)                                     | 17-20x 60-90 min<br>sessions (17-25.5<br>hours)  |
| Comparator  | Waitlist  | Parent training (CBT with parent only)                                   | Psychoeducational group. Students in both the treatment and comparison conditions received a tier 1 classroom-based psychoeducation and skills intervention, which was implemented throughout the school year  |
| Intervention length (weeks)                       | 20 umatic stress disorder: NR N   | 12   | 20   |

PTSD – Post-traumatic stress disorder; NR, Not relevant; DSM, Diagnostic and Statistical Manual of Mental Disorders; OCD, Obsessive Compulsive Disorder; ADHD, Attention Deficit Hyperactivity Disorder; TGCT, Trauma and Grief Component Therapy for Adolescents.

4 See appendix D for full evidence tables.

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#### 1 Quality assessment of clinical studies included in the evidence review

- 2 The clinical evidence profiles for this review (trauma-focused CBT for the treatment
- of PTSD in children and young people) are presented in Table 6, Table 7, Table 8,
- 4 Table 9, Table 10, Table 11, Table 12 and Table 13.

Table 6: Summary clinical evidence profile: Trauma-focused CBT versus meditation for early treatment (1-3 months)

Illustrative comparative risks\* (95% CI) Corresponding No of **Quality of Assumed** Relative **Participant** the risk Trauma-focused evidence effect **Outcomes** Meditation **CBT** (95% CI) (studies) (GRADE) **PTSD** The mean PTSD 31 low<sup>1</sup> symptomatology symptomatolo (1 study) clinician-rated at gy clinicianrated at 1-1-month followmonth followup in the intervention up CPTS-RI groups was change score 0.15 standard Follow-up: deviations lower mean 1 (0.85 lower to months 0.56 higher) **PTSD** The mean PTSD 30 low<sup>1</sup> symptomatolo symptomatology (1 study) gy clinicianclinician-rated at rated at 6-6-month followmonth followup in the intervention up CPTS-RI groups was change score 0.12 standard Follow-up: deviations higher mean 6 (0.6 lower to 0.83 months higher) Diagnosis at 333 per 1000 250 per 1000 RR 0.75 low1 31 1-month (83 to 760) (0.25 to (1 study) follow-up 2.28) Number of people who met criteria for a diagnosis of **PTSD** Follow-up: mean 1 months Diagnosis at 286 per 1000 189 per 1000 RR 0.66 30 low<sup>1</sup> (51 to 697) (1 study) 6-month (0.18 to follow-up 2.44)Number of people who met criteria for a diagnosis of **PTSD** 

|   | Illustrative co<br>(95% CI) | omparative risks*                     |                                |  |  |
|---|-----------------------------|---------------------------------------|--------------------------------|--|--|
| Outcomes  | Assumed risk Meditation     | Corresponding risk Trauma-focused CBT | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Follow-up:<br>mean 6<br>months  |                             |                                       |                                |  |  |
| Discontinuati on Number of participants lost to follow- up Follow-up: mean 1 months | 0                           | 0                                     | Not<br>estimabl<br>e           | 31<br>(1 study)                        | moderate <sup>2</sup>                    |

CBT=cognitive behavioural therapy; CI=confidence interval; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standard mean difference; CPTS-RI=Child Post-Traumatic Stress-Reaction Index.

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Table 7: Summary clinical evidence profile: Trauma-focused CBT versus waitlist. TAU or no treatment for delayed treatment (>3 months)

| waitii3t, 17   | Illustrative com                                    | _   | eu liea                       | minerit (~3                                | , months                                 |
|--|---|---|-------------------------------|--|--|
|  | risks* (95% CI)                                     |   | Rela<br>tive                  |  |  |
| Outcomes   | Assumed risk<br>Waitlist, TAU<br>or no<br>treatment | Correspon<br>ding risk<br>Trauma-<br>focused<br>CBT   | effe<br>ct<br>(95<br>%<br>CI) | No of<br>Particip<br>ants<br>(studie<br>s) | Quality of<br>the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatology<br>self-rated at<br>endpoint<br>SPTSS/CPSS/CRIE<br>S/CRTI/UCLA<br>PTSD-RI/CPTS-RI<br>change score<br>Follow-up: 0.4-13<br>weeks |   | The mean PTSD symptomato logy self-rated at endpoint in the intervention groups was 1.21 standard deviations lower (1.59 to 0.83 lower) |                               | 872<br>(13<br>studies)                     | very low <sup>1,2</sup>                  |
| PTSD symptomatology self-rated at 1-3 month follow-up IES/SPTSS/CRIES/ UCLA PTSD-RI/CPTS-RI change score Follow-up: 1-3 months                         |   | The mean PTSD symptomato logy self-rated at 1-3 month follow-up in the intervention groups was  |                               | 301<br>(5<br>studies)                      | low <sup>1,3</sup>                       |

<sup>&</sup>lt;sup>1</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm <sup>2</sup> OIS not met (events<300)

|  | 1.28<br>standard<br>deviations<br>lower<br>(1.68 to<br>0.87 lower)  |                       |                           |
|--|---|-----------------------|---------------------------|
| PTSD<br>symptomatology<br>self-rated at 6-<br>month follow-up<br>SPTSS change<br>score<br>Follow-up: mean 6<br>months        | The mean PTSD symptomato logy self-rated at 6-month follow-up in the intervention groups was 0.55 standard deviations lower (1.19 lower to 0.09 higher) | 39<br>(1<br>study)    | very low <sup>1,4</sup>   |
| PTSD<br>symptomatology<br>self-rated at 12-18<br>month follow-up<br>CPSS/SPTSS<br>change score<br>Follow-up: 12-18<br>months | The mean PTSD symptomato logy self-rated at 12-18 month follow-up in the intervention groups was 0.6 standard deviations lower (1.16 to 0.04 lower)     | 114<br>(2<br>studies) | very low <sup>1,3,5</sup> |
| PTSD symptomatology clinician-rated at endpoint CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score Follow-up: 8-20 weeks   | The mean PTSD symptomato logy clinician-rated at endpoint in the intervention groups was 1.47 standard deviations lower (2.03 to 0.9 lower)             | 409<br>(7<br>studies) | low <sup>5</sup>          |
| PTSD<br>symptomatology<br>clinician-rated at 3-<br>month follow-up   | The mean<br>PTSD<br>symptomato<br>logy  | 113<br>(3<br>studies) | very low <sup>1,3,6</sup> |

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| CAPS/K-SADS-E:<br>PTSD/ADIS-<br>C:PTSD change<br>score<br>Follow-up: mean 3<br>months  | clinician-<br>rated at 3-<br>month<br>follow-up in<br>the<br>intervention<br>groups was<br>0.75<br>standard<br>deviations<br>lower<br>(1.14 to<br>0.37 lower) |                      |                           |
|--|---|----------------------|---------------------------|
| PTSD symptomatology clinician-rated at 6-month follow-up CAPS/K-SADS-E: PTSD Follow-up: mean 6 months                                    | The mean PTSD symptomato logy clinician-rated at 6-month follow-up in the intervention groups was 0.69 standard deviations lower (1.12 to 0.25 lower)         | 89<br>(2<br>studies) | low <sup>3,6</sup>        |
| PTSD symptomatology clinician-rated at 12-month follow-up CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score Follow-up: mean 12 months | The mean PTSD symptomato logy clinician-rated at 12-month follow-up in the intervention groups was 0.63 standard deviations lower (1.09 to 0.16 lower)        | 88<br>(2<br>studies) | low <sup>3,6</sup>        |
| PTSD<br>symptomatology<br>clinician-rated at 2-<br>year follow-up<br>K-SADS-E: PTSD<br>change score<br>Follow-up: mean 2<br>years        | The mean PTSD symptomato logy clinician-rated at 2-year follow-up in the intervention groups was 0.22 standard  | 35<br>(1<br>study)   | very low <sup>1,4,6</sup> |

|   |              | da data   |   |                       |                           |
|---|--------------|---|---|-----------------------|---------------------------|
|   |              | deviations<br>lower<br>(0.9 lower to<br>0.46 higher)  |   |                       |                           |
| Remission at<br>endpoint<br>Number of people<br>no longer meeting<br>diagnostic criteria<br>for PTSD<br>Follow-up: 8-12<br>weeks  | 407 per 1000 | 712 per<br>1000<br>(541 to 936)   | RR<br>1.75<br>(1.33<br>to<br>2.3)       | 277<br>(5<br>studies) | moderate <sup>3</sup>     |
| Remission at 1-3<br>month follow-up<br>Number of people<br>no longer above<br>threshold on a scale<br>for PTSD or<br>meeting diagnostic<br>criteria for PTSD<br>Follow-up: 1-3<br>months  | 91 per 1000  | 666 per<br>1000<br>(258 to<br>1000)   | RR<br>7.33<br>(2.84<br>to<br>18.9<br>1) | 90<br>(2<br>studies)  | low <sup>6,7</sup>        |
| Remission at 12-18 month follow-up Number of people no longer meeting diagnostic criteria for PTSD/scoring above clinical threshold on a validated scale Follow-up: 12-18 months          | 324 per 1000 | 385 per<br>1000<br>(275 to 541)   | RR<br>1.19<br>(0.85<br>to<br>1.67)      | 213<br>(2<br>studies) | low <sup>4,6</sup>        |
| Response at endpoint Number of people showing clinically significant improvement, based on reliable change indices [RCI]/rated as 'much/very much improved' on CGI Follow-up: 10-13 weeks | 98 per 1000  | 525 per<br>1000<br>(161 to<br>1000)   | RR<br>5.35<br>(1.64<br>to<br>17.3<br>9) | 203<br>(3<br>studies) | very low <sup>1,5,7</sup> |
| Anxiety symptoms<br>at endpoint<br>HADS-<br>A/SCARED/RCMAS<br>/SCAS/BAI change<br>score<br>Follow-up: 2-20<br>weeks   |              | The mean anxiety symptoms at endpoint in the intervention groups was 0.81 standard deviations lower |   | 554<br>(8<br>studies) | very low <sup>1,5</sup>   |

|   | (1.23 to 0.4  |                        |                           |
|---|---|------------------------|---------------------------|
|   | lower)  |                        |                           |
| Anxiety symptoms<br>at 3-month follow-<br>up<br>HADS-A/RCMAS<br>change score<br>Follow-up: mean 3<br>months             | The mean anxiety symptoms at 3-month follow-up in the intervention groups was 0.34 standard deviations lower (1.18 lower to 0.5 higher) | 63<br>(2<br>studies)   | very low <sup>1,5,8</sup> |
| Anxiety symptoms<br>at 6-month follow-<br>up<br>HADS-A change<br>score<br>Follow-up: mean 6<br>months                   | The mean anxiety symptoms at 6-month follow-up in the intervention groups was 0.87 standard deviations lower (1.53 to 0.21 lower)       | 39<br>(1<br>study)     | very low <sup>1,3</sup>   |
| Anxiety symptoms<br>at 12-18 month<br>follow-up<br>HADS-A/SCARED<br>change score<br>Follow-up: 12-18<br>months          | The mean anxiety symptoms at 12-18 month follow-up in the intervention groups was 0.76 standard deviations lower (1.22 to 0.3 lower)    | 114<br>(2<br>studies)  | low <sup>1,3</sup>        |
| Depression<br>symptoms at<br>endpoint<br>HADS-D/CES-<br>D/CDI/MFQ/DSRS/<br>BDI change score<br>Follow-up: 2-20<br>weeks | The mean depression symptoms at endpoint in the intervention groups was 0.72 standard deviations lower                                  | 834<br>(13<br>studies) | low <sup>1,5</sup>        |

|  | (4.00.1   |                       |                           |
|--|---|-----------------------|---------------------------|
|  | (1.03 to<br>0.41 lower)   |                       |                           |
| Depression<br>symptoms at 1-3<br>month follow-up<br>BDI/HADS-D/CES-<br>D/CDI/MINI:Depres<br>sion /DSRS change<br>score<br>Follow-up: 1-3<br>months | The mean depression symptoms at 1-3 month follow-up in the intervention groups was 0.62 standard deviations lower (0.87 to 0.36 lower)  | 379<br>(7<br>studies) | very low <sup>1,3,6</sup> |
| Depression<br>symptoms at 6-<br>month follow-up<br>HADS-<br>D/CDI/MINI:Depres<br>sion change score<br>Follow-up: mean 6<br>months                  | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.48 standard deviations lower (0.84 to 0.13 lower)    | 129<br>(3<br>studies) | very low <sup>1,3,6</sup> |
| Depression<br>symptoms at 12-18<br>month follow-up<br>HADS-<br>D/CDI/MINI:Depres<br>sion/MFQ change<br>score<br>Follow-up: 12-18<br>months         | The mean depression symptoms at 12-18 month follow-up in the intervention groups was 0.5 standard deviations lower (0.78 to 0.22 lower) | 203<br>(4<br>studies) | low <sup>1,3</sup>        |
| Depression<br>symptoms at 2 year<br>follow-up<br>CDI change score<br>Follow-up: mean 2<br>years  | The mean depression symptoms at 2 year follow-up in the intervention groups was 0.17 standard deviations lower (0.83 lower              | 36<br>(1<br>study)    | very<br>low1,6,8          |

|  | to 0.5<br>higher)  |                       |                           |
|--|--|-----------------------|---------------------------|
| Emotional and<br>behavioural<br>problems at<br>endpoint<br>SDQ-A/CBCL<br>change score<br>Follow-up: 6-13<br>weeks  | The mean emotional and behavioural problems at endpoint in the intervention groups was 0.58 standard deviations lower (0.79 to 0.36 lower)                       | 476<br>(5<br>studies) | low <sup>1</sup>          |
| Emotional and<br>behavioural<br>problems at 18-<br>month follow-up<br>SDQ change score<br>Follow-up: mean 18<br>months                                   | The mean emotional and behavioural problems at 18-month follow-up in the intervention groups was 2.83 lower (4.79 to 0.87 lower)                                 | 75<br>(1<br>study)    | low <sup>1,3</sup>        |
| Emotional and behavioural problems- Externalizing at endpoint CBCL Externalizing change score Follow-up: 12-20 weeks                                     | The mean emotional and behavioural problems-externalizin g at endpoint in the intervention groups was 0.25 standard deviations lower (0.67 lower to 0.16 higher) | 210<br>(3<br>studies) | very low <sup>1,4</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at 3-<br>month follow-up<br>CBCL Externalizing<br>change score<br>Follow-up: mean 3<br>months | The mean emotional and behavioural problems-externalizin g at 3-month follow-up in the   | 56<br>(2<br>studies)  | very low <sup>1,3,6</sup> |

|  | intervention<br>groups was<br>0.77<br>standard<br>deviations<br>lower<br>(1.32 to<br>0.21 lower)  |                    |                           |
|--|---|--------------------|---------------------------|
| Emotional and behavioural problems- Externalizing at 6- month follow-up CBCL Externalizing change score Follow-up: mean 6 months                           | The mean emotional and behavioural problems-externalizin g at 6-month follow-up in the intervention groups was 0.82 standard deviations lower (1.57 to 0.07 lower)        | 32<br>(1<br>study) | very low <sup>1,3,6</sup> |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at 12-<br>month follow-up<br>CBCL Externalizing<br>change score<br>Follow-up: mean 12<br>months | The mean emotional and behavioural problems-externalizin g at 12-month follow-up in the intervention groups was 0.7 standard deviations lower (1.44 lower to 0.04 higher) | 32<br>(1<br>study) | very low <sup>1,4,6</sup> |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at 2-<br>year follow-up<br>CBCL Externalizing<br>change score<br>Follow-up: mean 2<br>years     | The mean emotional and behavioural problems-externalizin g at 2-year follow-up in the intervention groups was 1.41 standard deviations lower                              | 32<br>(1<br>study) | very low <sup>1,3,6</sup> |

|  | (2.22 to<br>0.61 lower)  |                       |                           |
|--|--|-----------------------|---------------------------|
| Emotional and<br>behavioural<br>problems-<br>Internalizing at<br>endpoint<br>CBCL Internalizing<br>change score<br>Follow-up: 12-20<br>weeks             | The mean emotional and behavioural problems-internalizing at endpoint in the intervention groups was 0.61 standard deviations lower (1.03 to 0.2 lower)                  | 178<br>(2<br>studies) | very low <sup>1,3</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Internalizing at 3-<br>month follow-up<br>CBCL Internalizing<br>change score<br>Follow-up: mean 3<br>months | The mean emotional and behavioural problems-internalizing at 3-month follow-up in the intervention groups was 0.71 standard deviations lower (1.54 lower to 0.12 higher) | 24<br>(1<br>study)    | very low <sup>1,4,6</sup> |
| Quality of life<br>KIDSCREEN-27:<br>Global HRQoL T-<br>scores/ILK; change<br>score<br>Follow-up: 6-12<br>weeks<br>Better indicated by<br>higher values   | The mean quality of life in the intervention groups was 0.33 standard deviations higher (0.06 to 0.6 higher)   | 219<br>(2<br>studies) | very low <sup>1,3</sup>   |
| Functional impairment at endpoint CAPS: Functional impairment/SAS-SR-Y change score Follow-up: 10-13 weeks   | The mean functional impairment at endpoint in the intervention groups was 1.56 standard deviations lower   | 95<br>(2<br>studies)  | very low <sup>1,2,4</sup> |

|  | (0.44.1  |                       |                                |
|--|--|-----------------------|--------------------------------|
|  | (3.14 lower<br>to 0.02<br>higher)  |                       |                                |
| Functional impairment at 3-month follow-up CAPS: Functional impairment; change score Follow-up: mean 3 months                            | The mean functional impairment at 3-month follow-up in the intervention groups was 0.96 standard deviations lower (1.24 to 0.68 lower)       | 220<br>(2<br>studies) | very<br>low <sup>1,3,5,6</sup> |
| Functional impairment at 6-month follow-up CAPS: Functional impairment; change score Follow-up: mean 6 months                            | The mean functional impairment at 6-month follow-up in the intervention groups was 0.45 standard deviations lower (0.99 lower to 0.1 higher) | 54<br>(1<br>study)    | very low <sup>1,4,6</sup>      |
| Functional impairment at 12-month follow-up CAPS: Functional impairment; change score Follow-up: mean 12 months                          | The mean functional impairment at 12-month follow-up in the intervention groups was 1.28 standard deviations lower (1.88 to 0.69 lower)      | 53<br>(1<br>study)    | very low <sup>1,3,6</sup>      |
| Global functioning<br>at endpoint<br>CGAS/fCPSS/GAF<br>change score<br>Follow-up: 10-20<br>weeks<br>Better indicated by<br>higher values | The mean global functioning at endpoint in the intervention groups was 1.25 standard deviations higher                                       | 321<br>(4<br>studies) | very low <sup>1,3,5</sup>      |

|  |             | (0.65 to<br>1.85 higher)  |                                   |                         |                           |
|--|-------------|---|-----------------------------------|-------------------------|---------------------------|
| Global functioning<br>at 3-month follow-<br>up<br>GAF; change score<br>Follow-up: mean 3<br>months<br>Better indicated by<br>higher values       |             | The mean global functioning at 3-month follow-up in the intervention groups was 1.35 standard deviations higher (0.45 to 2.25 higher)       |                                   | 24<br>(1<br>study)      | very low <sup>1,3,6</sup> |
| Global functioning<br>at 18-month follow-<br>up<br>fCPSS change<br>score<br>Follow-up: mean 18<br>months<br>Better indicated by<br>higher values |             | The mean global functioning at 18-month follow-up in the intervention groups was 0.1 standard deviations higher (0.35 lower to 0.56 higher) |                                   | 75<br>(1<br>study)      | low <sup>1,4</sup>        |
| Discontinuation<br>Number of<br>participants lost to<br>follow-up for any<br>reason<br>Follow-up: 0.4-20<br>weeks                                | 75 per 1000 | 98 per 1000<br>(70 to 137)  | RR<br>1.3<br>(0.93<br>to<br>1.83) | 1255<br>(18<br>studies) | moderate <sup>9</sup>     |

ADIS-C=Anxiety Disorder Interview Schedule-Child version: BAI=Beck Anxiety Index; BDI=Beck Depression Inventory; CAPS=Clinician Administered PTSD Symptom; CBCL=Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI=Children's Depression Inventory; CES-D=Centre for Epidemiological Studies-Depression; CGAS= Children's Global Assessment Scale; CGI=Clinical Global Impression, CI=confidence interval; CPSS=Child PTSD Symptom Scale; CPTS-RI=Child Post-Traumatic Symptom-Reaction Index; CRIES=Children's Revised Impact of Event Scale; CRTI=Children's Response to Trauma Inventory; DSRS=Depression Self-Rating Scale; GAF=Global Assessment of Functioning; HRQoL=Health-Related Quality of Life; KIDSCREEN-27=Health-related quality of life questionnaire for children, young people and their parents; K-SADS-E=Kiddie Schedule for Affective Disorders and Schizophrenia-Epidemiological; HADS-A/D=Hospital Anxiety and Depression Scale-Anxiety/Depression; ILK=an instrument to measure quality of life in children and adolescents; MFQ=Mood and Feeling Questionnaire; PTSD=post-traumatic stress disorder; RCMAS=Revised Children's Manifest Anxiety Scale; RR=risk ratio; SAS-SR=Social Adjustment Scale-Self Report; SCARED=Screen for Child Anxiety Related Disorders; SCAS=Spence Children's Anxiety Scale; SDQ =Strength and Difficulties Questionnaires; SMD=standard mean difference; SPTSS=Screen for Post-Traumatic Stress Symptoms; TAU=treatment as usual; UCLA PTSD-RI=UCLA PTSD-Reaction Index <sup>1</sup> Risk of bias is high or unclear across multiple domains

1234567890112345678 1112345678 <sup>2</sup> Considerable heterogeneity (I2>80%)

<sup>3</sup> OIS not met (N<400)

- <sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit
- 19 20 21 <sup>5</sup> Substantial heterogeneity (I2=>50%)
  - <sup>6</sup> Data is not reported/cannot be extracted for all outcomes
    - <sup>7</sup> OIS not met (events<300)

1

3

4

 $^8$  95% CI crosses line of no effect and thresholds for both clinically important benefit and harm  $^9$  95% CI crosses both line of no effect and threshold for clinically important harm

Table 8: Summary clinical evidence profile: Trauma-focused CBT versus supportive counselling for delayed treatment (>3 months)

| Supportive   | Illustrative co                     | r delayed treatr   | nent (>3 n                     | ionthis                                | <b>6</b> III                   |
|--|-------------------------------------|--|--------------------------------|--|--------------------------------|
|  | risks* (95% CI                      |  |                                |  | Quality of the                 |
| Outcomes   | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE<br>)    |
| PTSD<br>symptomatology self-<br>rated at endpoint<br>CRIES/TSCC-<br>PTSD/UCLA PTSD-<br>RI/CPSS change<br>score<br>Follow-up: 6-15<br>weeks |                                     | The mean PTSD symptomatolog y self-rated at endpoint in the intervention groups was 0.49 standard deviations lower (0.71 to 0.26 lower)          |                                | 325<br>(5<br>studies)                  | low <sup>1,2</sup>             |
| PTSD<br>symptomatology self-<br>rated at 3-month<br>follow-up<br>CRIES change score<br>Follow-up: mean 3<br>months                         |                                     | The mean PTSD symptomatolog y self-rated at 3-month follow-up in the intervention groups was 1.58 standard deviations lower (2.62 to 0.55 lower) |                                | 20<br>(1 study)                        | very<br>low <sup>1,2</sup>     |
| PTSD<br>symptomatology self-<br>rated at 6-month<br>follow-up<br>TSCC-PTSD/VCPSS<br>change score<br>Follow-up: mean 6<br>months            |                                     | The mean PTSD symptomatolog y self-rated at 6-month follow-up in the intervention groups was 0.7 standard deviations lower (1.29 to 0.11 lower)  |                                | 120<br>(2<br>studies)                  | very<br>low <sup>1,2,3,4</sup> |
| PTSD<br>symptomatology self-<br>rated at 12-17 month<br>follow-up<br>TSCC-PTSD/CPSS<br>change score<br>Follow-up: 12-17<br>months          |                                     | The mean PSTD symptomatolog y self-rated at 12-17 month follow-up in the intervention groups was   |                                | 181<br>(3<br>studies)                  | very<br>low <sup>1,2,4</sup>   |

|   | Illustrative co                     | mnarative  |                                |  | 0                           |
|---|-------------------------------------|--|--------------------------------|--|-----------------------------|
|   | risks* (95% CI                      |  |                                |  | Quality of the              |
| Outcomes  | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE<br>) |
|   | <b>.</b>                            | 0.69 standard<br>deviations<br>lower<br>(0.99 to 0.39<br>lower)  | (,                             |  | ,                           |
| PTSD<br>symptomatology<br>clinician-rated at<br>endpoint<br>K-SADS-PL:<br>PTSD/CPSS-<br>I/CAPS; change<br>score<br>Follow-up: 8-14<br>weeks |                                     | The mean PTSD symptomatolog y clinician-rated at endpoint in the intervention groups was 0.71 standard deviations lower (1.1 to 0.31 lower)                  |                                | 231<br>(3<br>studies)                  | moderat<br>e <sup>2</sup>   |
| PTSD<br>symptomatology<br>clinician-rated at 3-<br>month follow-up<br>CAPS change score<br>Follow-up: mean 3<br>months                      |                                     | The mean PTSD symptomatolog y clinician-rated at 3-month follow-up in the intervention groups was 0.25 standard deviations lower (0.81 lower to 0.31 higher) |                                | 50<br>(1 study)                        | low <sup>4,5</sup>          |
| PTSD<br>symptomatology<br>clinician-rated at 6-<br>month follow-up<br>CAPS change score<br>Follow-up: mean 6<br>months                      |                                     | The mean PTSD symptomatolog y clinician-rated at 6-month follow-up in the intervention groups was 0.43 standard deviations lower (1 lower to 0.13 higher)    |                                | 49<br>(1 study)                        | low <sup>4,5</sup>          |
| PTSD<br>symptomatology<br>clinician-rated at 12-<br>month follow-up<br>CAPS/CPSS-I<br>change score  |                                     | The mean PTSD symptomatolog y clinician- rated at 12- month follow-  |                                | 109<br>(2<br>studies)                  | low <sup>2,4</sup>          |

|  | Illustrative co  | · ·   |                                |  | Quality                   |
|--|--|---|--------------------------------|--|---------------------------|
| Outcomes   | risks* (95% CI<br>Assumed<br>risk<br>Supportive<br>counselling | Correspondin g risk Trauma- focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | of the evidenc e (GRADE   |
| Follow-up: mean 12 months  | Counciling   | up in the intervention groups was 0.89 standard deviations lower (1.28 to 0.49 lower) | (66% 61)                       | (Studies)                              | ,                         |
| Remission at endpoint Number of people no longer meeting diagnostic criteria for PTSD Follow-up: 8-15 weeks  | 376 per 1000   | 628 per 1000<br>(470 to 839)  | RR 1.67<br>(1.25 to<br>2.23)   | 208<br>(4<br>studies)                  | moderat<br>e <sup>6</sup> |
| Remission at 6-month follow-up Number of people no longer meeting diagnostic criteria for PTSD Follow-up: mean 6 months                                      | 263 per 1000   | 632 per 1000<br>(276 to 1000)   | RR 2.4<br>(1.05 to<br>5.49)    | 38<br>(1 study)                        | moderat<br>e <sup>6</sup> |
| Remission at 12-<br>month follow-up<br>Number of people no<br>longer meeting<br>diagnostic criteria for<br>PTSD<br>Follow-up: mean 12<br>months              | 500 per 1000   | 780 per 1000<br>(585 to 1000)   | RR 1.56<br>(1.17 to<br>2.08)   | 118<br>(2<br>studies)                  | low <sup>4,6</sup>        |
| Response at endpoint Number of people showing clinically significant improvement (based on RCI) Follow-up: mean 14 weeks                                     | 267 per 1000   | 741 per 1000<br>(395 to 1000)   | RR 2.78<br>(1.48 to<br>5.22)   | 61<br>(1 study)                        | low <sup>4,6</sup>        |
| Response at 12-<br>month follow-up<br>Number of people<br>showing clinically<br>significant<br>improvement (based<br>on RCI)<br>Follow-up: mean 12<br>months | 400 per 1000   | 708 per 1000<br>(432 to 1000)   | RR 1.77<br>(1.08 to<br>2.9)    | 61<br>(1 study)                        | low <sup>4,6</sup>        |

|   | Illustrative corrisks* (95% CI      |   |                                |  | Quality of the                 |
|---|-------------------------------------|---|--------------------------------|--|--------------------------------|
| Outcomes  | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE         |
| Dissociative<br>symptoms at<br>endpoint<br>TSCC-Dissociation<br>change score<br>Follow-up: mean 12<br>weeks             |                                     | The mean dissociative symptoms at endpoint in the intervention groups was 0.27 standard deviations lower (0.71 lower to 0.16 higher)    |                                | 82<br>(1 study)                        | very<br>low <sup>1,4,5</sup>   |
| Dissociative<br>symptoms at 6-<br>month follow-up<br>TSCC-Dissociation<br>change score<br>Follow-up: mean 6<br>months   |                                     | The mean dissociative symptoms at 6-month follow-up in the intervention groups was 0.7 standard deviations lower (1.15 to 0.25 lower)   |                                | 82<br>(1 study)                        | very<br>low <sup>1,2,4</sup>   |
| Dissociative<br>symptoms at 12-<br>month follow-up<br>TSCC-Dissociation<br>change score<br>Follow-up: mean 12<br>months |                                     | The mean dissociative symptoms at 12-month follow-up in the intervention groups was 0.49 standard deviations lower (0.93 to 0.05 lower) |                                | 82<br>(1 study)                        | very<br>low <sup>1,2,4</sup>   |
| Anxiety symptoms at endpoint STAI-State/SCARED/TSC C:Anxiety change score Follow-up: 8-12 weeks                         |                                     | The mean anxiety symptoms at endpoint in the intervention groups was 0.29 standard deviations lower (0.48 to 0.1 lower)                 |                                | 433<br>(4<br>studies)                  | low <sup>1,4</sup>             |
| Anxiety symptoms at<br>6-month follow-up<br>STAI-State change<br>score  |                                     | The mean anxiety symptoms at 6-month follow-up in the   |                                | 233<br>(2<br>studies)                  | very<br>low <sup>1,3,4,5</sup> |

|   | Illustrative co                                 |   |                 |                           | Quality                        |
|---|---|---|-----------------|---------------------------|--------------------------------|
|   | risks* (95% CI<br>Assumed<br>risk<br>Supportive | Correspondin<br>g risk<br>Trauma-   | Relative effect | No of<br>Participa<br>nts | of the evidenc e (GRADE        |
| Outcomes Follow-up: mean 6 months   | counselling                                     | intervention<br>groups was<br>0.3 standard<br>deviations<br>lower<br>(0.82 lower to<br>0.22 higher)   | (95% CI)        | (studies)                 | )                              |
| Anxiety symptoms at<br>12-month follow-up<br>STAI-State change<br>score<br>Follow-up: mean 12<br>months                     |   | The mean anxiety symptoms at 12-month follow-up in the intervention groups was 0.17 standard deviations lower (0.51 lower to 0.17 higher)   |                 | 237<br>(2<br>studies)     | very<br>low <sup>1,4,5</sup>   |
| Depression<br>symptoms at<br>endpoint<br>BDI/CES-<br>D/CDI/TSCC:Depres<br>sion change score<br>Follow-up: 6-15<br>weeks     |   | The mean depression symptoms at endpoint in the intervention groups was 0.41 standard deviations lower (0.67 to 0.16 lower)                 |                 | 552<br>(7<br>studies)     | low <sup>1,4</sup>             |
| Depression<br>symptoms at 3-<br>month follow-up<br>CES-<br>D/MINI:Depression<br>change score<br>Follow-up: mean 3<br>months |   | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.46 standard deviations lower (2.26 lower to 1.33 higher) |                 | 70<br>(2<br>studies)      | very<br>low <sup>1,4,7,8</sup> |
| Depression<br>symptoms at 6-<br>month follow-up<br>BDI/CDI/MINI:Depre<br>ssion change score<br>Follow-up: mean 6<br>months  |   | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.3 standard deviations lower                              |                 | 320<br>(4<br>studies)     | very<br>low <sup>1,3,4,5</sup> |

|  | Illustrative corrisks* (95% CI      | •   |                                |  | Quality of the                 |
|--|-------------------------------------|---|--------------------------------|--|--------------------------------|
| Outcomes   | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE         |
|  |                                     | (0.74 lower to 0.13 higher)   | ,                              |  |                                |
| Depression<br>symptoms at 12-17<br>month follow-up<br>BDI/CDI/MINI:Depre<br>ssion change score<br>Follow-up: 12-17<br>months                               |                                     | The mean depression symptoms at 12-17 month follow-up in the intervention groups was 0.34 standard deviations lower (0.74 lower to 0.07 higher)                           |                                | 384<br>(5<br>studies)                  | very<br>low <sup>1,3,4,5</sup> |
| Emotional and<br>behavioural<br>problems-<br>Internalizing at<br>endpoint<br>CBCL Internalizing<br>change score<br>Follow-up: mean 12<br>weeks             |                                     | The mean emotional and behavioural problems-internalizing at endpoint in the intervention groups was 0.08 standard deviations lower (0.33 lower to 0.16 higher)           |                                | 261<br>(2<br>studies)                  | very<br>low <sup>1,2,4</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Internalizing at 6-<br>month follow-up<br>CBCL Internalizing<br>change score<br>Follow-up: mean 6<br>months   |                                     | The mean emotional and behavioural problems-internalizing at 6-month follow-up in the intervention groups was 0.17 standard deviations higher (0.19 lower to 0.53 higher) |                                | 224<br>(2<br>studies)                  | very<br>low <sup>1,4,9</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Internalizing at 12-<br>month follow-up<br>CBCL Internalizing<br>change score<br>Follow-up: mean 12<br>months |                                     | The mean emotional and behavioural problems-internalizing at 12-month follow-up in the intervention groups was 0.02 standard deviations                                   |                                | 228<br>(2<br>studies)                  | very<br>low <sup>1,2,4</sup>   |

|  | Illustrative comparative                        |  |                 |                       | Quality                        |
|--|---|--|-----------------|-----------------------|--------------------------------|
| Outcomes   | risks* (95% CI<br>Assumed<br>risk<br>Supportive | Correspondin<br>g risk<br>Trauma-  | Relative effect | No of Participa nts   | of the evidence (GRADE         |
| Outcomes   | counselling                                     | higher<br>(0.24 lower to<br>0.28 higher)   | (95% CI)        | (studies)             | )                              |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at<br>endpoint<br>CBCL Externalizing<br>change score<br>Follow-up: mean 12<br>months            |   | The mean emotional and behavioural problems-externalizing at endpoint in the intervention groups was 0.15 standard deviations lower (0.4 lower to 0.09 higher)             |                 | 261<br>(2<br>studies) | very<br>low <sup>1,2,4</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at 6-<br>month follow-up<br>CBCL Externalizing<br>change score<br>Follow-up: mean 6<br>months   |   | The mean emotional and behavioural problems-externalizing at 6-month follow-up in the intervention groups was 0.04 standard deviations higher (0.22 lower to 0.31 higher)  |                 | 224<br>(2<br>studies) | very<br>low <sup>1,2,4</sup>   |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at 12-<br>month follow-up<br>CBCL Externalizing<br>change score<br>Follow-up: mean 12<br>months |   | The mean emotional and behavioural problems-externalizing at 12-month follow-up in the intervention groups was 0.18 standard deviations higher (0.27 lower to 0.62 higher) |                 | 228<br>(2<br>studies) | very<br>low <sup>1,3,4,9</sup> |
| Behaviour problems<br>at endpoint<br>CBCL total score;<br>change score<br>Follow-up: 8-12<br>weeks   |   | The mean behaviour problems at endpoint in the intervention groups was 0.11 standard deviations  |                 | 385<br>(3<br>studies) | very<br>low <sup>1,2,4</sup>   |

|  |                                     | Ilustrative comparative  |                                |  | Quality of the               |
|--|-------------------------------------|--|--------------------------------|--|------------------------------|
| Outcomes   | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE       |
|  | <b>J</b>                            | lower<br>(0.31 lower to<br>0.09 higher)  |                                | (                                      | ,                            |
| Behaviour problems<br>at 6-month follow-up<br>CBCL total score;<br>change score<br>Follow-up: mean 6<br>months       |                                     | The mean behaviour problems at 6-month follow-up in the intervention groups was 0.08 standard deviations higher (0.18 lower to 0.34 higher)    |                                | 224<br>(2<br>studies)                  | very<br>low <sup>1,2,4</sup> |
| Behaviour problems<br>at 12-month follow-<br>up<br>CBCL total score;<br>change score<br>Follow-up: mean 12<br>months |                                     | The mean behaviour problems at 12-month follow-up in the intervention groups was 0.04 standard deviations higher (0.32 lower to 0.41 higher)   |                                | 228<br>(2<br>studies)                  | very<br>low <sup>1,2,4</sup> |
| Functional impairment at 3-month follow-up CAPS: Functional impairment; change score Follow-up: mean 3 months        |                                     | The mean functional impairment at 3-month follow-up in the intervention groups was 0.43 standard deviations lower (1 lower to 0.13 higher)     |                                | 50<br>(1 study)                        | very<br>low <sup>1,4,5</sup> |
| Functional impairment at 6-month follow-up CAPS: Functional impairment; change score Follow-up: mean 6 months        |                                     | The mean functional impairment at 6-month follow-up in the intervention groups was 0.01 standard deviations higher (0.55 lower to 0.57 higher) |                                | 49<br>(1 study)                        | very<br>low <sup>1,4,8</sup> |

|   | Illustrative co                     |  |                                |  | Quality of the               |
|---|-------------------------------------|--|--------------------------------|--|------------------------------|
| Outcomes  | Assumed risk Supportive counselling | Correspondin<br>g risk<br>Trauma-<br>focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE       |
| Functional impairment at 12-month follow-up CAPS: Functional impairment; change score Follow-up: mean 12 months                           |                                     | The mean functional impairment at 12-month follow-up in the intervention groups was 1.12 standard deviations lower (1.73 to 0.5 lower) |                                | 48<br>(1 study)                        | very<br>low <sup>1,2,4</sup> |
| Global functioning at endpoint CGAS; change score Follow-up: 14-15 weeks Better indicated by higher values                                |                                     | The mean global functioning at endpoint in the intervention groups was 1.08 standard deviations higher (0.65 to 1.5 higher)            |                                | 99<br>(2<br>studies)                   | very<br>low <sup>1,2,4</sup> |
| Global functioning at<br>6-month follow-up<br>CGAS; change score<br>Follow-up: mean 6<br>months<br>Better indicated by<br>higher values   |                                     | The mean global functioning at 6-month follow-up in the intervention groups was 1.05 standard deviations higher (0.37 to 1.73 higher)  |                                | 38<br>(1 study)                        | low <sup>1,2</sup>           |
| Global functioning at<br>12-month follow-up<br>CGAS; change score<br>Follow-up: mean 12<br>months<br>Better indicated by<br>higher values |                                     | The mean global functioning at 12-month follow-up in the intervention groups was 1 standard deviations higher (0.47 to 1.54 higher)    |                                | 61<br>(1 study)                        | very<br>low <sup>1,2,4</sup> |
| Discontinuation<br>Number of<br>participants lost to<br>follow-up for any   | 287 per 1000                        | 224 per 1000<br>(175 to 290)   | RR 0.78<br>(0.61 to<br>1.01)   | 678<br>(8<br>studies)                  | moderat<br>e <sup>5</sup>    |

|                                    | Illustrative comparative risks* (95% CI) |  |                                |  | Quality of the              |
|------------------------------------|--|--|--------------------------------|--|-----------------------------|
| Outcomes                           | Assumed risk Supportive counselling      | Correspondin<br>g risk<br>Trauma-<br>focused CBT | Relative<br>effect<br>(95% CI) | No of<br>Participa<br>nts<br>(studies) | evidenc<br>e<br>(GRADE<br>) |
| reason<br>Follow-up: 3-15<br>weeks |  |  |                                |  |                             |

BDI= Beck Depression Inventory; CAPS= Clinician Administered PTSD Symptom; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CES-D= Centre for Epidemiological Studies-Depression; CGAS= Children's Global Assessment Scale; CI=confidence interval; CPSS= Child PTSD Symptom Scale; CRIES= Children's Revised Impact of Event Scale; K-SADS= Kiddie Schedule for Affective Disorders and Schizophrenia-Epidemiological; PTSD=post-traumatic stress disorder; RCI=Reliable Change Indecies; RR=risk ratio; SCARED=Screen for Child Anxiety Related Disorders; SMD=standardised mean difference; STAI=State-Trait Anxiety Inventory; TSCC =Trauma Symptom Checklist for Children; UCLA PTSD-RI=UCLA PTSD-Reaction Index

- 1 2 3 4 5 6 7 8 9 10 <sup>1</sup> Risk of bias is high or unclear across multiple domains
  - <sup>2</sup> OIS not met (N<400)
- 11 12 13 14 15 <sup>3</sup> Substantial heterogeneity (I2>50%)
  - <sup>4</sup> Data is not reported/cannot be extracted for all outcomes
  - 5 95% CI crosses both line of no effect and threshold for clinically important benefit
    - <sup>6</sup> OIS not met (events<300)

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- 16 17 <sup>7</sup> Considerable heterogeneity (I2>80%)
  - 8 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm
  - <sup>9</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

Table 9: Summary clinical evidence profile: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for delayed treatment (>3 months)

| ti outii  | ient (>5 months  |   |                                       |  |   |
|---|--|---|---------------------------------------|--|---|
|   | Illustrative com   | parative risks* (95%  |                                       |  |   |
| Outcomes  | Assumed risk Eye movement desensitisatio n and reprocessing (EMDR) | Corresponding risk Trauma-focused CBT   | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |
| PTSD<br>symptomatolo<br>gy self-rated<br>at endpoint<br>CRTI change<br>score<br>Follow-up:<br>mean 6<br>weeks |  | The mean PTSD simptomatology self-rated at endpoint in the intervention groups was 0.13 standard deviations lower (0.56 lower to 0.29 higher) |                                       | 85<br>(1 study)                        | very<br>low <sup>1,2,3</sup>                |
| PTSD<br>symptomatolo<br>gy self-rated<br>at 3-month<br>follow-up<br>CRTI change<br>score                      |  | The mean PTSD symptomatology self-rated at 3-month follow-up in the intervention groups was 0.35 standard                                     |                                       | 85<br>(1 study)                        | very<br>low <sup>1,2,3</sup>                |

|  | Illustrative comp  | parative risks* (95%  |                                       |  |                                   |
|--|--|---|---------------------------------------|--|-----------------------------------|
| Outcomes   | Assumed risk Eye movement desensitisatio n and reprocessing (EMDR) | Corresponding risk Trauma-focused CBT   | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality of the evidenc e (GRADE ) |
| Follow-up:<br>mean 3<br>months   |  | deviations lower<br>(0.77 lower to 0.08<br>higher)  |                                       |  |                                   |
| PTSD symptomatolo gy self-rated at 12-month follow-up CRTI change score Follow-up: mean 12 months                                |  | The mean PTSD symptomatology self-rated at 12-month follow-up in the intervention groups was 0.24 standard deviations lower (0.66 lower to 0.19 higher) |                                       | 85<br>(1 study)                        | very<br>low <sup>1,2,3</sup>      |
| PTSD<br>symptomatolo<br>gy clinician-<br>rated<br>CAPS-CA<br>change score<br>Follow-up:<br>mean 8<br>weeks                       |  | The mean PTSD symptomatology clinician-rated in the intervention groups was 0.04 standard deviations higher (0.53 lower to 0.6 higher)                  |                                       | 48<br>(1 study)                        | very<br>low <sup>3,4</sup>        |
| Emotional and<br>behavioural<br>problems at<br>endpoint<br>SDQ-A<br>change score<br>Follow-up:<br>mean 6<br>weeks                |  | The mean emotional and behavioural problems at endpoint in the intervention groups was 0.55 standard deviations higher (0.12 to 0.99 higher)            |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,5</sup>      |
| Emotional and<br>behavioural<br>problems at 3-<br>month follow-<br>up<br>SDQ-A<br>change score<br>Follow-up:<br>mean 3<br>months |  | The mean emotional and behavioural problems at 3-month follow-up in the intervention groups was 0.46 standard deviations higher (0.03 to 0.89 higher)   |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,5</sup>      |
| Emotional and<br>behavioural<br>problems at<br>12-month  |  | The mean emotional and behavioural problems at 12-  |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,5</sup>      |

|   | Illustrative comp  | parative risks* (95%  |                                       |  |   |
|---|--|---|---------------------------------------|--|---|
| Outcomes  | Assumed risk Eye movement desensitisatio n and reprocessing (EMDR) | Corresponding risk Trauma-focused CBT   | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |
| follow-up<br>SDQ-A<br>change score<br>Follow-up:<br>mean 12<br>months   |  | month follow-up in<br>the intervention<br>groups was<br>0.45 standard<br>deviations higher<br>(0.02 to 0.89<br>higher)                  |                                       |  |   |
| Quality of life<br>at endpoint<br>KIDSCREEN-<br>27: Global<br>HRQoL T-<br>scores;<br>change score<br>Follow-up:<br>mean 6<br>weeks<br>Better<br>indicated by<br>higher values |  | The mean quality of life at endpoint in the intervention groups was 0.23 standard deviations lower (0.66 lower to 0.2 higher)           |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,6</sup>                |
| Quality of life at 3-month follow-up KIDSCREEN-27: Global HRQoL T-scores; change score Follow-up: mean 3 months Better indicated by higher values                             |  | The mean quality of life at 3-month follow-up in the intervention groups was 0.39 standard deviations lower (0.82 lower to 0.04 higher) |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,6</sup>                |
| Quality of life at 12-month follow-up KIDSCREEN-27: Global HRQoL T-scores; change score Follow-up: mean 12 months Better indicated by higher values                           |  | The mean quality of life at 12-month follow-up in the intervention groups was 0.3 standard deviations lower (0.73 lower to 0.12 higher) |                                       | 85<br>(1 study)                        | very<br>low <sup>1,3,6</sup>                |

|  | Illustrative comparative risks* (95% CI)                           |                                       |                                       |  |   |
|--|--|---------------------------------------|---------------------------------------|--|---|
| Outcomes   | Assumed risk Eye movement desensitisatio n and reprocessing (EMDR) | Corresponding risk Trauma-focused CBT | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |
| Discontinuation Number of participants lost to follow-up for any reason Follow-up: 6-8 weeks | 118 per 1000   | 94 per 1000<br>(36 to 241)            | RR 0.8<br>(0.31<br>to<br>2.05)        | 133<br>(2 studies)                     | low <sup>4</sup>                            |

CAPS=Clinician Administered PTSD Symptom;; CBT=cognitive behavioural therapy; Cl=confidence interval; CRTI= Children's Response to Trauma Inventory; EMDR=Eye Movement Desensitisation and Reprocessing; HRQoL=Health-Related Quality of Life; KIDSCREEN-27= Health-related quality of life questionnaire for children, young people and their parents; PTSD=post-traumatic stress disorder; RR=risk ratio; SDQ-A= Strength and Difficulties Questionnaires; SMD=standard mean difference.

<sup>1</sup> Risk of bias is high or unclear across multiple domains

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Table 10: Summary clinical evidence profile: Trauma-focused CBT versus combined somatic and cognitive therapies for delayed treatment (>3 months)

|   | -,  |  |                                |  |  |
|---|---|--|--------------------------------|--|--|
|   | Illustrative c<br>(95% CI)                            | omparative risks*  |                                |  |  |
| Outcomes  | Assumed risk Combined somatic and cognitive therapies | Corresponding risk Trauma-focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatolo<br>gy self-rated<br>at endpoint<br>SPTSS<br>change score<br>Follow-up:<br>mean 2 weeks |   | The mean PTSD symptomatology self-rated at endpoint in the intervention groups was 0.87 standard deviations higher (0.21 to 1.53 higher) |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| PTSD<br>symptomatolo<br>gy self-rated<br>at 3-month<br>follow-up<br>SPTSS                                   |   | The mean PTSD symptomatology self-rated at 3-month follow-up in the intervention groups was  |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>5</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>6</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

|   | Illustrative c  | omparative risks*  |                                |  |  |
|---|---|--|--------------------------------|--|--|
| Outcomes  | Assumed risk Combined somatic and cognitive therapies | Corresponding<br>risk<br>Trauma-focused<br>CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| change score<br>Follow-up:<br>mean 3<br>months  |   | 0.8 standard<br>deviations higher<br>(0.15 to 1.46<br>higher)  |                                |  |  |
| PTSD<br>symptomatolo<br>gy self-rated<br>at 6-month<br>follow-up<br>SPTSS<br>change score<br>Follow-up:<br>mean 6<br>months   |   | The mean PTSD symptomatology self-rated at 6-month follow-up in the intervention groups was 0.83 standard deviations higher (0.17 to 1.48 higher)  |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| PTSD<br>symptomatolo<br>gy self-rated<br>at 12-month<br>follow-up<br>SPTSS<br>change score<br>Follow-up:<br>mean 12<br>months |   | The mean PTSD symptomatology self-rated at 12-month follow-up in the intervention groups was 0.92 standard deviations higher (0.26 to 1.58 higher) |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>endpoint<br>HADS-A<br>change score<br>Follow-up:<br>mean 2 weeks                                    |   | The mean anxiety symptoms at endpoint in the intervention groups was 1.01 standard deviations higher (0.34 to 1.68 higher)                         |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>3-month<br>follow-up<br>HADS-A<br>change score<br>Follow-up:<br>mean 3<br>months                    |   | The mean anxiety symptoms at 3-month follow-up in the intervention groups was 0.91 standard deviations higher (0.25 to 1.57 higher)                |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>6-month<br>follow-up<br>HADS-A<br>change score  |   | The mean anxiety symptoms at 6-month follow-up in the intervention groups was 0.22 standard  |                                | 39<br>(1 study)                        | very<br>low <sup>1,3</sup>               |

|   | Illustrative c  | omparative risks*  |                                |  |  |
|---|---|--|--------------------------------|--|--|
| Outcomes  | Assumed risk Combined somatic and cognitive therapies | Corresponding risk Trauma-focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Follow-up:<br>mean 6<br>months  |   | deviations higher<br>(0.41 lower to 0.85<br>higher)  |                                |  |  |
| Anxiety symptoms at 12-month follow-up HADS-A change score Follow-up: mean 12 months                          |   | The mean anxiety symptoms at 12-month follow-up in the intervention groups was 0.09 standard deviations lower (0.71 lower to 0.54 higher)    |                                | 39<br>(1 study)                        | very<br>low <sup>1,4</sup>               |
| Depression<br>symptoms at<br>endpoint<br>HADS-D<br>change score<br>Follow-up:<br>mean 2 weeks                 |   | The mean depression symptoms at endpoint in the intervention groups was 1.3 standard deviations higher (0.6 to 1.99 higher)                  |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Depression<br>symptoms at<br>3-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:<br>mean 3<br>months |   | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.45 standard deviations higher (0.19 lower to 1.09 higher) |                                | 39<br>(1 study)                        | very<br>low <sup>1,3</sup>               |
| Depression<br>symptoms at<br>6-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:<br>mean 6<br>months |   | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.3 standard deviations higher (0.33 lower to 0.93 higher)  |                                | 39<br>(1 study)                        | very<br>low <sup>1,3</sup>               |
| Depression<br>symptoms at<br>12-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:                    |   | The mean depression symptoms at 12-month follow-up in the intervention groups was 0.66 standard  |                                | 39<br>(1 study)                        | very<br>low <sup>1,2</sup>               |

|  | Illustrative c<br>(95% CI)                            | omparative risks*                             |                                |  |  |
|--|---|---|--------------------------------|--|--|
| Outcomes   | Assumed risk Combined somatic and cognitive therapies | Corresponding risk Trauma-focused CBT         | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| mean 12<br>months  |   | deviations higher<br>(0.02 to 1.31<br>higher) | , ,                            |  | Ì  |
| Discontinuation Number of participants lost to follow- up for any reason Follow-up: mean 2 weeks | 0 per 1000  | 0 per 1000<br>(0 to 0)                        | RR 3<br>(0.13 to<br>69.52)     | 40<br>(1 study)                        | very<br>low <sup>1,4</sup>               |

CBT=cognitive behavioural therapy; CI=confidence interval; HADS-A/D= Hospital Anxiety and Depression Scale-Anxiety/Depression; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; SPTSS= Screen for Post-Traumatic Stress Symptoms

<sup>1</sup> Risk of bias is high or unclear across multiple domains

1234567

Table 11: Summary clinical evidence profile: Trauma-focused CBT + parent training versus waitlist for delayed treatment (>3 months)

|   | Illustrative<br>(95% CI) | comparative risks*   | Ì                                  | ·                                      |  |
|---|--------------------------|--|------------------------------------|--|--|
| Outcomes  | Assumed risk Waitlist    | Corresponding risk Trauma-focused CBT + parent training  | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participan<br>ts<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatolog<br>y clinician-rated<br>at endpoint<br>ADIS-C: PTSD;<br>change score<br>Follow-up:<br>mean 20 weeks             |                          | The mean PTSD symptomatology clinician-rated at endpoint in the intervention groups was 1.73 standard deviations lower (2.69 to 0.77 lower)          |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |
| PTSD<br>symptomatolog<br>y clinician-rated<br>at 3-month<br>follow-up<br>ADIS-C: PTSD;<br>change score<br>Follow-up:<br>mean 3 months |                          | The mean PTSD symptomatology clinician-rated at 3-month follow-up in the intervention groups was 1.34 standard deviations lower (2.24 to 0.44 lower) |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

|   | Illustrative<br>(95% CI) | comparative risks*  |                                    |  |  |
|---|--------------------------|---|------------------------------------|--|--|
| Outcomes  | Assumed risk Waitlist    | Corresponding risk Trauma-focused CBT + parent training   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participan<br>ts<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Anxiety<br>symptoms at<br>endpoint<br>RCMAS;<br>change score<br>Follow-up:<br>mean 20 weeks   |                          | The mean anxiety symptoms at endpoint in the intervention groups was 0.33 standard deviations lower (1.13 lower to 0.48 higher)                                 |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |
| Anxiety<br>symptoms at 3-<br>month follow-up<br>RCMAS;<br>change score<br>Follow-up:<br>mean 3 months   |                          | The mean anxiety symptoms at 3-month follow-up in the intervention groups was 0.75 standard deviations lower (1.58 lower to 0.09 higher)                        |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |
| Depression<br>symptoms at<br>endpoint<br>CDI; change<br>score<br>Follow-up:<br>mean 20 weeks  |                          | The mean depression symptoms at endpoint in the intervention groups was 0.61 standard deviations lower (1.43 lower to 0.21 higher)                              |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |
| Depression<br>symptoms at 3-<br>month follow-up<br>CDI; change<br>score<br>Follow-up:<br>mean 3 months  |                          | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.36 standard deviations lower (1.17 lower to 0.45 higher)                     |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |
| Emotional and<br>behavioural<br>problems-<br>Internalizing at<br>endpoint<br>CBCL:<br>Internalizing;<br>change score<br>Follow-up:<br>mean 20 weeks |                          | The mean emotional and behavioural problems-internalizing at endpoint in the intervention groups was 0.45 standard deviations lower (1.26 lower to 0.36 higher) |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |

|  | Illustrative<br>(95% CI) | comparative risks*  |                                    |  |  |
|--|--------------------------|---|------------------------------------|--|--|
| Outcomes   | Assumed risk Waitlist    | Corresponding risk Trauma-focused CBT + parent training   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participan<br>ts<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Emotional and behavioural problems-Internalizing at 3-month follow-up CBCL: Internalizing; change score Follow-up: mean 3 months                                 |                          | The mean emotional and behavioural problems-internalizing at 3-month follow-up in the intervention groups was 0.92 standard deviations lower (1.77 to 0.07 lower) |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at<br>endpoint<br>CBCL:<br>Externalizing;<br>change score<br>Follow-up:<br>mean 20 weeks              |                          | The mean emotional and behavioural problems-externalizing at endpoint in the intervention groups was 0.44 standard deviations lower (1.25 lower to 0.37 higher)   |                                    | 24<br>(1 study)                        | very low <sup>1,3,4</sup>                |
| Emotional and<br>behavioural<br>problems-<br>Externalizing at<br>3-month follow-<br>up<br>CBCL:<br>Externalizing;<br>change score<br>Follow-up:<br>mean 3 months |                          | The mean emotional and behavioural problems-externalizing at 3-month follow-up in the intervention groups was 0.88 standard deviations lower (1.73 to 0.04 lower) |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |
| Global<br>functioning at<br>endpoint<br>GAF; change<br>score<br>Follow-up:<br>mean 20 weeks<br>Better indicated<br>by higher<br>values                           |                          | The mean global functioning at endpoint in the intervention groups was 2.02 standard deviations higher (1.01 to 3.04 higher)                                      |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |
| Global<br>functioning at 3-<br>month follow-up<br>GAF; change<br>score   |                          | The mean global functioning at 3-month follow-up in the intervention groups was   |                                    | 24<br>(1 study)                        | very low <sup>1,2,3</sup>                |

|  | Illustrative comparative risks* (95% CI) |  |                                    |  |  |
|--|--|--|------------------------------------|--|--|
| Outcomes   | Assumed risk Waitlist                    | Corresponding risk Trauma-focused CBT + parent training        | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participan<br>ts<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Follow-up:<br>mean 3 months<br>Better indicated<br>by higher<br>values   |  | 2.04 standard<br>deviations higher<br>(1.02 to 3.06<br>higher) |                                    |  |  |
| Discontinuation<br>Number of<br>participants lost<br>to follow-up for<br>any reason<br>Follow-up:<br>mean 20 weeks | 167 per<br>1000                          | 250 per 1000<br>(50 to 1000)                                   | RR 1.5<br>(0.3 to<br>7.43)         | 24<br>(1 study)                        | low <sup>5</sup>                         |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBT=cognitive behavioural therapy; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS= Revised Children's Manifest Anxiety Scale; RR=risk ratio; SMD=standardised mean difference

12345678910

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Table 12: Summary clinical evidence profile: Trauma-focused CBT versus parent training (CBT with parent-only) for delayed treatment (>3 months)

|  | Illustrative co<br>(95% CI)                         | omparative risks*  |                                |                                    |  |
|--|---|--|--------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk Parent training (CBT with parent-only) | Corresponding risk Trauma-focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD symptomatolo gy clinician-rated at endpoint K-SADS-E: PTSD; change score Follow-up: mean 12 weeks |   | The mean PTSD symptomatology clinician-rated at endpoint in the intervention groups was 0.34 standard deviations lower (0.96 lower to 0.27 higher) |                                | 41<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| PTSD<br>symptomatolo<br>gy clinician-<br>rated at 3-<br>month follow-                                  |   | The mean PTSD symptomatology clinician-rated at 3-month follow-up in the intervention  |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>5</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

|   | Illustrative comparative risks* (95% CI)            |   |                                |                                    |  |
|---|---|---|--------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk Parent training (CBT with parent-only) | Corresponding risk Trauma-focused CBT   | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| up K-SADS-E: PTSD; change score Follow-up: mean 3 months  |   | groups was<br>0.12 standard<br>deviations higher<br>(0.49 lower to 0.73<br>higher)  |                                |                                    |  |
| PTSD symptomatolo gy clinician-rated at 6-month follow-up K-SADS-E: PTSD; change score Follow-up: mean 6 months   |   | The mean PTSD symptomatology clinician-rated at 6-month follow-up in the intervention groups was 0.25 standard deviations lower (0.87 lower to 0.36 higher)   |                                | 41<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| PTSD symptomatolo gy clinician-rated at 12-month follow-up K-SADS-E: PTSD; change score Follow-up: mean 12 months |   | The mean PTSD symptomatology clinician-rated at 12-month follow-up in the intervention groups was 0.07 standard deviations higher (0.54 lower to 0.68 higher) |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |
| PTSD symptomatolo gy clinician-rated at 2-year follow-up K-SADS-E: PTSD; change score Follow-up: mean 2 years     |   | The mean PTSD symptomatology clinician-rated at 2-year follow-up in the intervention groups was 0.64 standard deviations higher (0.01 to 1.27 higher)         |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,6</sup>             |
| Emotional and behavioural problems- Externalizing at endpoint CBCL Externalizing                                  |   | The mean emotional and behavioural problems-externalizing at endpoint in the intervention groups  |                                | 38<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |

|   | Illustrative comparative risks* (95% CI)            |  |                                |                                    |  |
|---|---|--|--------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk Parent training (CBT with parent-only) | Corresponding risk Trauma-focused CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| change score<br>Follow-up:<br>mean 12<br>weeks  |   | was 0.13 standard deviations higher (0.51 lower to 0.77 higher)  |                                |                                    |  |
| Emotional and behavioural problems- Externalizing at 3-month follow-up CBCL Externalizing change score Follow-up: mean 3 months   |   | The mean emotional and behavioural problems-externalizing at 3-month follow-up in the intervention groups was 0.61 standard deviations lower (1.27 lower to 0.04 higher) |                                | 38<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Emotional and behavioural problems- Externalizing at 6-month follow-up CBCL Externalizing change score Follow-up: mean 6 months   |   | The mean emotional and behavioural problems-externalizing at 6-month follow-up in the intervention groups was 0.75 standard deviations lower (1.41 to 0.09 lower)        |                                | 38<br>(1 study)                    | very<br>low <sup>1,3,6</sup>             |
| Emotional and behavioural problems- Externalizing at 12-month follow-up CBCL Externalizing change score Follow-up: mean 12 months |   | The mean emotional and behavioural problems-externalizing at 12-month follow-up in the intervention groups was 0.79 standard deviations lower (1.45 to 0.12 lower)       |                                | 38<br>(1 study)                    | very<br>low <sup>1,3,6</sup>             |
| Emotional and<br>behavioural<br>problems-<br>Externalizing<br>at 2-year<br>follow-up<br>CBCL                                      |   | The mean emotional and behavioural problems-externalizing at 2-year follow-up in the intervention  |                                | 38<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |

|  | Illustrative comparative risks* (95% CI)            |   |                                |                                    |  |
|--|---|---|--------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk Parent training (CBT with parent-only) | Corresponding<br>risk<br>Trauma-focused<br>CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Externalizing change score Follow-up: mean 2 years   |   | groups was<br>0.53 standard<br>deviations lower<br>(1.18 lower to 0.12<br>higher)   |                                |                                    |  |
| Depression<br>symptoms at<br>endpoint<br>CDI change<br>score<br>Follow-up:<br>mean 12<br>weeks               |   | The mean depression symptoms at endpoint in the intervention groups was 0.3 standard deviations higher (0.32 lower to 0.92 higher)            |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression<br>symptoms at<br>3-month<br>follow-up<br>CDI change<br>score<br>Follow-up:<br>mean 3<br>months   |   | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.12 standard deviations higher (0.49 lower to 0.73 higher)  |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression<br>symptoms at<br>6-month<br>follow-up<br>CDI change<br>score<br>Follow-up:<br>mean 6<br>months   |   | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.09 standard deviations higher (0.53 lower to 0.7 higher)   |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |
| Depression<br>symptoms at<br>12-month<br>follow-up<br>CDI change<br>score<br>Follow-up:<br>mean 12<br>months |   | The mean depression symptoms at 12-month follow-up in the intervention groups was 0.31 standard deviations higher (0.31 lower to 0.93 higher) |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression symptoms at 2-year follow-  |   | The mean depression symptoms at 2-  |                                | 41<br>(1 study)                    | very<br>low <sup>1,3,6</sup>             |

|   | Illustrative comparative risks* (95% CI)            |   |                                |                                    |  |
|---|---|---|--------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk Parent training (CBT with parent-only) | Corresponding<br>risk<br>Trauma-focused<br>CBT  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| up<br>CDI change<br>score<br>Follow-up:<br>mean 2 years |   | year follow-up in<br>the intervention<br>groups was<br>0.73 standard<br>deviations higher<br>(0.1 to 1.37 higher) |                                |                                    |  |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS=; RR=risk ratio; SMD=standardised mean difference

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Table 13: Summary clinical evidence profile: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for delayed treatment (>3 months)

| delayed freatment (>5 months)   |                                       |   |                                    |  |   |  |  |
|---|---------------------------------------|---|------------------------------------|--|---|--|--|
|   | Illustrative com<br>(95% CI)          | Illustrative comparative risks* (95% CI)  |                                    |  |   |  |  |
| Outcomes  | Assumed risk Psychoeducat ional group | Corresponding risk Trauma-focused CBT (+ psychoeducation al group)  | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |  |  |
| PTSD<br>symptomatolo<br>gy self-rated<br>at endpoint<br>UCLA PTSD-<br>RI; change<br>score<br>Follow-up:<br>mean 20<br>weeks |                                       | The mean PTSD symptomatology self-rated at endpoint in the intervention groups was 0.46 standard deviations lower (0.81 to 0.11 lower)          |                                    | 127<br>(1 study)                       | low <sup>1,2</sup>                          |  |  |
| PTSD symptomatolo gy self-rated at 4 month follow-up UCLA PTSD-RI; change score Follow-up: mean 4 months                    |                                       | The mean PTSD symptomatology self-rated at 4 month follow-up in the intervention groups was 0.57 standard deviations lower (1.07 to 0.07 lower) |                                    | 65<br>(1 study)                        | low <sup>1,2</sup>                          |  |  |

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple outcomes

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

 $<sup>^{\</sup>rm 5}$  95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>6</sup> OIS not met (N<400)

|  | Illustrative comparative risks* (95% CI) |  |                                    |  |   |
|--|--|--|------------------------------------|--|---|
| Outcomes   | Assumed risk Psychoeducat ional group    | Corresponding risk Trauma-focused CBT (+ psychoeducation al group)   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |
| Response at endpoint Number of people showing clinically significant improvement, based on reliable change indices (RCI) Follow-up: mean 20 weeks          | 244 per 1000                             | 493 per 1000<br>(317 to 768)   | RR 2.02<br>(1.3 to<br>3.15)        | 159<br>(1 study)                       | low <sup>1,3</sup>                          |
| Response at 4-month follow-up Number of people showing clinically significant improvement, based on reliable change indices (RCI) Follow-up: mean 4 months | 171 per 1000                             | 377 per 1000<br>(215 to 657)   | RR 2.21<br>(1.26 to<br>3.85)       | 159<br>(1 study)                       | low <sup>1,3</sup>                          |
| Depression<br>symptoms at<br>endpoint<br>DSRS<br>change score<br>Follow-up:<br>mean 20<br>weeks  |  | The mean depression symptoms at endpoint in the intervention groups was 0.44 standard deviations lower (0.8 to 0.09 lower)           |                                    | 125<br>(1 study)                       | low <sup>1,2</sup>                          |
| Depression<br>symptoms at<br>4 month<br>follow-up<br>DSRS<br>change score<br>Follow-up:<br>mean 4<br>months  |  | The mean depression symptoms at 4 month follow-up in the intervention groups was 0.59 standard deviations lower (1.08 to 0.09 lower) |                                    | 66<br>(1 study)                        | low <sup>1,2</sup>                          |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

|   | Illustrative comparative risks* (95% CI) |  |                                    |  |   |
|---|--|--|------------------------------------|--|---|
| Outcomes  | Assumed risk Psychoeducat ional group    | Corresponding risk Trauma-focused CBT (+ psychoeducation al group) | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidenc<br>e<br>(GRADE |
| Discontinuation Number of participants lost to follow- up for any reason Follow-up: mean 20 weeks | 256 per 1000                             | 143 per 1000<br>(74 to 277)  | RR 0.56<br>(0.29 to<br>1.08)       | 159<br>(1 study)                       | moderate<br>4                               |

- CBT=cognitive behavioural therapy; CI=confidence interval; DSRS= Depression Self-Rating Scale;
- 1234567 PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standard mean difference; UCLA PTSD-
- RI=UCLA PTSD-Reaction Index
- <sup>1</sup> Risk of bias is high or unclear across multiple outcomes
- <sup>2</sup> OIS not met (N<400)
- <sup>3</sup> OIS not met (events<300)
- <sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm
- 8 See appendix F for full GRADE tables.

## 9 Sensitivity and subgroup analysis

- 10 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- 11 treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- 12 by multiplicity of trauma revealed a statistically significant subgroup difference for
- clinician-rated PTSD symptomatology (Chi<sup>2</sup> = 15.60, p < 0.0001), with a relatively 13
- larger effect observed for single incident index trauma (SMD -2.80 [-3.62, -1.99]) 14
- 15 compared with multiple incident index trauma (SMD -0.98 [-1.37, -0.59]), although
- 16 both effects are large and statistically significant. Non-significant subgroup
- 17 differences were observed for self-rated PTSD symptomatology and discontinuation.
- 18 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- 19 treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- by specific intervention revealed a statistically significant subgroup difference for 20
- clinician-rated PTSD symptomatology (Chi² = 28.74, p < 0.00001), with relatively 21
- 22 larger effects observed for cognitive therapy (SMD -2.80 [-3.62, -1.99]) and narrative
- 23 exposure therapy (SMD -1.87 [-2.84, -0.90]), although effects were clinically
- 24 important and statistically significant across all specific interventions. Non-significant
- 25 subgroup differences were observed for self-rated PTSD symptomatology and
- 26 discontinuation.
- 27 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- 28 treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- 29 by format revealed a statistically significant difference for self-rated PTSD
- symptomatology (Chi<sup>2</sup> = 8.52, p = 0.04), with relatively larger effects observed for 30
- individual (child-only; SMD -1.82 [-2.43, -1.21]) and caregiver and child (SMD -1.25 [-31
- 32 2.09, -0.42]) compared with group (SMD -0.72 [-1.16, -0.28]), although effects were
- clinically important and statistically significant across all formats. Non-significant 33

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- 1 subgroup differences were observed for clinician-rated PTSD symptomatology and
- 2 discontinuation.
- 3 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- 4 treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- 5 by age range revealed non-significant subgroup differences on PTSD
- 6 symptomatology (self-rated and clinician-rated) and discontinuation, between studies
- 7 where the age range includes children aged 7 years and under and studies where
- 8 the age range only includes over 7s.
- 9 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- by diagnostic status at baseline revealed a statistically significant subgroup
- difference for clinician-rated PTSD symptomatology (Chi² = 4.58, p = 0.03), with
- 13 relatively larger effects observed for those with a diagnosis at baseline (SMD -2.31 [-
- 14 3.26, -1.36]) compared to those with clinically important PTSD symptoms (scoring
- above a threshold on a validated scale) but not necessarily a diagnosis at baseline,
- although effects are large and statistically significant for both subgroups. Non-
- 17 significant subgroup differences were observed for self-rated PTSD symptomatology
- 18 and discontinuation.
- 19 Sub-analysis of the comparison trauma-focused CBT versus waitlist, TAU or no
- treatment for delayed treatment (>3 months) of clinically important symptoms/PTSD,
- 21 by trauma type revealed a statistically significant subgroup difference for clinician-
- rated PTSD symptomatology (Chi<sup>2</sup> = 28.74, p < 0.00001), with relatively larger effects
- observed for children exposed to motor vehicle collisions (SMD -2.80 [-3.62, -1.99])
- or witnessing war as a civilian (SMD -1.87 [-2.84, -0.90]), although effects are
- 25 clinically important and statistically significant across trauma types. Non-significant
- 26 subgroup differences were observed for self-rated PTSD symptomatology and
- 27 discontinuation.
- 28 Sub-analysis of the comparison trauma-focused CBT versus supportive counselling
- 29 for delayed treatment (>3 months) of clinically important symptoms/PTSD, by
- 30 multiplicity of trauma revealed non-significant subgroup differences for self-rated
- 31 PTSD symptomatology and discontinuation, and sub-analysis was not possible for
- 32 clinician-rated PTSD as there is only a single subgroup (multiple incident index
- 33 trauma).
- 34 Sub-analysis of the comparison trauma-focused CBT versus supportive counselling
- for delayed treatment (>3 months) of clinically important symptoms/PTSD by specific
- intervention, by format, by age range, by diagnostic status at baseline, and by trauma
- 37 type revealed non-significant subgroup differences for PTSD symptomatology (self-
- rated and clinician-rated) and discontinuation.
- 39 See forest plots in Appendix E.

## 40 Non-trauma-focused cognitive behavioural therapies (CBT): clinical

### 41 evidence

#### 42 Included studies

- 43 Five studies of non-trauma-focused CBT for the treatment of PTSD in children and
- 44 young people were identified for full-text review. Of these 3 studies, 1 RCT (N=33)
- was included in a single comparison for non-trauma-focused CBT.

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- 1 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 2 symptoms, there were no included studies.
- 3 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 4 PTSD symptoms, 1 RCT (N=33) compared non-trauma-focused CBT in addition to
- 5 TAU with TAU-only (Najavits 2006)

#### 6 Excluded studies

- 7 Four studies were reviewed at full text and excluded from this review because the
- 8 intervention was not targeted at PTSD symptoms, group assignment was non-
- 9 randomised, or the paper was a systematic review with no new useable data and any
- meta-analysis results not appropriate to extract.
- 11 Studies not included in this review with reasons for their exclusions are provided in
- 12 Appendix K.

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## 13 Summary of clinical studies included in the evidence review

- 14 Table 14 provides a brief summary of the included study and evidence from this
- 15 study is summarised in the clinical GRADE evidence profile below (Table 15).
- 16 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 17 study evidence tables in Appendix D.

## Table 14: Summary of included studies: Non-trauma-focused CBT for delayed treatment (>3 months)

| treatment (> 3 months)                   |  |  |  |  |
|--|--|--|--|--|
| Comparison                               | Non-trauma focused CBT (+ TAU) versus TAU  |  |  |  |
| Total no. of studies (N randomised)      | 1 (33)   |  |  |  |
| Study ID                                 | Najavits 2006  |  |  |  |
| Country                                  | US   |  |  |  |
| Diagnostic status                        | PTSD diagnosis according to ICD/DSM criteria   |  |  |  |
| Mean months since onset of PTSD          | 61.2   |  |  |  |
| Mean age (range)                         | 16.1 (range NR)  |  |  |  |
| Sex (% female)                           | 100  |  |  |  |
| Ethnicity (% BME)                        | 21   |  |  |  |
| Coexisting conditions                    | All met current DSM-IV criteria for both PTSD and SUD, with 94% having substance dependence. Current substance dependence diagnoses per DSM-IV criteria at intake were: cannabis (79%), alcohol (67%), hallucinogens (21%), amphetamines (15%), cocaine (9%), opioids (9%), inhalants (9%), barbiturates (6%), polysubstance (6%), and PCP 1 (3%). Participants could have more than one diagnosis |  |  |  |
| Mean months since traumatic event        | 88 (average age of first trauma was 8.75)  |  |  |  |
| Type of traumatic event                  | Mixed: The most common trauma category was sexual abuse (88%), followed by general disaster/accident (82%), physical abuse (73%), and crime (39%)  |  |  |  |
| Single or multiple incident index trauma | Multiple   |  |  |  |

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| Comparison                    | Non-trauma focused CBT (+ TAU) versus TAU  |
|-------------------------------|--|
| Lifetime experience of trauma | NR   |
| Intervention details          | Seeking Safety (based on manual by Najavits 2002) + TAU  |
| Intervention format           | Individual   |
| Intervention intensity        | 25x 50-min sessions (20.8 hours) + 1 session with carer.  Mean attended seeking safety sessions 9.7 (5.1) (+ 1.33 [SD = 2.09] sessions of trauma discussion; 0.78 sessions [SD = 1.00] of unspecified therapy) |
| Comparator                    | TAU: All participants were allowed to attend any treatments they naturalistically sought (e.g., Alcoholics Anonymous, psychotropic medication, and other individual and group psychotherapies)                 |
| Intervention length (weeks)   | 13   |

1 NR-Not reported; TAU-Treatment as usual.

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- 2 See appendix D for full evidence tables.
- 3 Quality assessment of clinical studies included in the evidence review
- 4 The clinical evidence profile for this review (non-trauma-focused CBT for the
- 5 treatment of PTSD in children and young people) is presented in Table 15.

Table 15: Summary clinical evidence profile: Non-trauma focused CBT (+ TAU) versus TAU for delayed treatment (>3 months)

|   | Illustrative comparative risks* (95% CI) |   | ,                              |                                    |  |
|---|--|---|--------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk                             | Corresponding risk Non-trauma focused CBT (+TAU)  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Depression<br>symptoms at<br>endpoint<br>Adolescent<br>Psychopatholog<br>y Scale: Axis I -<br>Major<br>Depression;<br>change score<br>Follow-up:<br>mean 13 weeks           |  | The mean depression symptoms at endpoint in the intervention groups was 0.33 standard deviations lower (1.02 lower to 0.37 higher)  |                                | 33<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Depression<br>symptoms at 3-<br>month follow-up<br>Adolescent<br>Psychopatholog<br>y Scale: Axis I -<br>Major<br>Depression;<br>change score<br>Follow-up:<br>mean 3 months |  | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.71 standard deviations higher (0 to 1.42 higher) |                                | 33<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |

|  | Illustrative comparative risks* (95% CI) |  |                                |                                    |  |
|--|--|--|--------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk                             | Corresponding risk Non-trauma focused CBT (+TAU)   | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Substance use disorder symptoms at endpoint Adolescent Psychopatholog y Scale: Axis I - Substance Use Disorder; change score Follow-up: mean 13 weeks          |  | The mean substance use disorder symptoms at endpoint in the intervention groups was 1.03 standard deviations lower (1.77 to 0.3 lower)                   |                                | 33<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |
| Substance use disorder symptoms at 3-month follow-up Adolescent Psychopatholog y Scale: Axis I - Substance Use Disorder; change score Follow-up: mean 3 months |  | The mean substance use disorder symptoms at 3-month follow-up in the intervention groups was 0.63 standard deviations higher (0.08 lower to 1.33 higher) |                                | 33<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Discontinuation<br>Number of<br>participants lost<br>to follow-up for<br>any reason<br>Follow-up:<br>mean 13 weeks   | 200 per<br>1000                          | 222 per 1000<br>(58 to 842)  | RR 1.11<br>(0.29 to<br>4.21)   | 33<br>(1 study)                    | low <sup>6</sup>                         |

- 12345678 CBT=cognitive behavioural therapy; CI=confidence interval; PTSD=post-traumatic stress disorder;
- RR=risk ratio; SMD=standardised mean difference; TAU=treatment as usual
  - <sup>1</sup> Risk of bias is high or unclear across multiple outcomes
- <sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit
- <sup>3</sup> Data is not reported/cannot be extracted for all outcomes
  - <sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm
- <sup>5</sup> OIS not met (N<400)
- <sup>6</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm
- 9 See appendix F for full GRADE tables.

## 10 Behavioural therapies: clinical evidence

## 11 Included studies

- 12 Three studies of behavioural therapies for the treatment of PTSD in children and
- 13 young people were identified for full-text review. None of these studies could not be
- 14 included.

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### 1 Excluded studies

- 2 Three studies were reviewed at full text and excluded from this review because the
- 3 intervention was not targeted at PTSD symptoms, the paper was a systematic review
- 4 with no new useable data and any meta-analysis results not appropriate to extract, or
- 5 the reference was a book section.
- 6 Studies not included in this review with reasons for their exclusions are provided in
- 7 Appendix K.

## 8 Psychologically-focused debriefing: clinical evidence

#### 9 Included studies

- Two studies of psychologically-focused debriefing for the treatment of PTSD in
- 11 children and young people were identified for full-text review. Neither of these studies
- 12 could be included.

#### 13 Excluded studies

- 14 Two studies were reviewed at full text and excluded from this review due to non-
- randomised group assignment or because the paper was a commentary.
- 16 Studies not included in this review with reasons for their exclusions are provided in
- 17 Appendix K.

## 18 Eye movement desensitisation and reprocessing (EMDR): clinical evidence

## 19 Included studies

- 20 Eleven studies of eye movement desensitisation and reprocessing (EMDR) for the
- 21 treatment of PTSD in children and young people were identified for full-text review.
- 22 Of these 11 studies, 3 RCTs (N=165) were included in a single comparison for
- 23 EMDR.
- 24 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- symptoms, there were no included studies.
- 26 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 27 PTSD symptoms, 3 RCTs (N=165) compared EMDR with waitlist or TAU (Ahmad
- 28 2007/ Ahmad & Sundelin-Wahlsten 2008 [one study reported across two papers]; de
- 29 Roos 2017; Soberman 2002).
- 30 Comparison with trauma-focused CBT are presented in the Trauma-focused CBT
- 31 section above.

#### 32 Excluded studies

- 33 Eight studies were reviewed at full text and excluded from this review. The most
- 34 common reasons for exclusion were that the intervention was not targeted at PTSD
- 35 symptoms or the paper was a systematic review with no new useable data and any
- meta-analysis results not appropriate to extract.
- 37 Studies not included in this review with reasons for their exclusions are provided in
- 38 Appendix K.

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Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## 1 Summary of clinical studies included in the evidence review

- See also the study selection flow chart in Appendix C, forest plots in Appendix E and study evidence tables in Appendix D.
- Table 16 provides brief summaries of the included studies and evidence from these are summarised in the clinical GRADE evidence profile below (Table 17).
- See also the study selection flow chart in Appendix C, forest plots in Appendix E and study evidence tables in Appendix D.

## Table 16: Summary of included studies: Eye movement desensitisation and reprocessing (EMDR) for delayed treatment (>3 months)

| reprocessing (EMDR) for delayed treatment (>3 months) |   |  |  |  |  |  |
|---|---|--|--|--|--|--|
| Comparison  | EMDR versus waitlist or TAU   |  |  |  |  |  |
| Total no. of studies (N randomised)                   | 3 (165)   |  |  |  |  |  |
| Study ID  | Ahmad 2007/2008 <sup>1</sup><br>de Roos 2017 <sup>2</sup><br>Soberman 2002 <sup>3</sup>   |  |  |  |  |  |
| Country   | Sweden <sup>1</sup> Netherlands <sup>2</sup> US <sup>3</sup>  |  |  |  |  |  |
| Diagnostic status                                     | PTSD diagnosis according to ICD/DSM criteria <sup>1</sup> Clinically important PTSD symptoms (scoring above a threshold on validated scale) <sup>2,3</sup>  |  |  |  |  |  |
| Mean months since onset of PTSD                       | Mean NR (the duration between the traumatic event and the establishment of a PTSD diagnosis was less than 1 year for 18.2% of the subjects, 1-2 years for 48.5%, and more than 3 years for 33.3%) <sup>1</sup> NR <sup>2,3</sup>  |  |  |  |  |  |
| Mean age (range)                                      | 9.9 (6-16) <sup>1</sup><br>13.1(8-18) <sup>2</sup><br>Mean NR (10-16) <sup>3</sup>  |  |  |  |  |  |
| Sex (% female)  | 61 <sup>1</sup><br>57 <sup>2</sup><br>0 <sup>3</sup>  |  |  |  |  |  |
| Ethnicity (% BME)                                     | NR  |  |  |  |  |  |
| Coexisting conditions                                 | 79% fulfilled DSM-IV criteria for at least one additional diagnosis: Depression (46%); ADHD (30%); ODD (21%); separation anxiety (18%); conduct disorder (12%), overanxious disorder and autism spectrum (3%) <sup>1</sup> 54% had one or more co-morbid disorder (assessed with ADIS-C) <sup>2</sup> |  |  |  |  |  |
|   | Other primary diagnoses included: Conduct Disorder (59%);<br>Attention Deficit Hyperactive Disorder (17%), Learning<br>Disability (14%), Substance Abuse (13%), and<br>Oppositional/Defiant Disorder (3%) <sup>3</sup>  |  |  |  |  |  |
| Mean months since traumatic event                     | Mean NR (the age when experiencing trauma was below 6 years in 33.3% of subjects, 7-10 years in 45.5%, and above 11 years in 21.2%) <sup>1</sup> 16.5 <sup>2</sup> NR <sup>3</sup>  |  |  |  |  |  |

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| Comparison                               | EMDR versus waitlist or TAU   |  |  |  |
|--|---|--|--|--|
| Type of traumatic event                  | Mixed: Maltreatment (36.4%), sexual abuse (21.2%), road accident (15.2%), witnessing unnatural death (12.1%) and other types of trauma (6.1%) <sup>1</sup> Mixed: Physical abuse/assault (23%); Sexual abuse (26%); Accident/injury of a loved one (19%); Traumatic loss (18%); Disaster/other (13%) <sup>2</sup> Unclear (no details reported) <sup>3</sup>              |  |  |  |
| Single or multiple incident index trauma | Multiple <sup>1</sup> Single <sup>2</sup> Unclear <sup>3</sup>  |  |  |  |
| Lifetime experience of trauma            | NR  |  |  |  |
| Intervention details                     | EMDR protocol (Shapiro 1995) adjusted for child age and developmental level <sup>1</sup>  |  |  |  |
|  | Eye movement desensitisation and reprocessing (EMDR) based on the standard protocol from Shapiro (2001) with age-appropriate modifications suggested by Tinker and Wilson (1999) and Greenwald (1999) <sup>2</sup>  |  |  |  |
|  | Eye movement desensitisation and reprocessing (EMDR, following the manual by Shapiro 1995 with selected population-specific variations suggested by Greenwald 1999), in addition to the usual treatment in either a residential or day treatment program at the same facility <sup>3</sup>  |  |  |  |
| Intervention format                      | Individual  |  |  |  |
| Intervention intensity                   | 8x weekly 45-min sessions (6 hours). Mean number of session provided 5.9 (range 1-8). 59% completed at least 7/8 sessions and 82% at least 4/8 sessions <sup>1</sup> 6x weekly 45-min sessions (4.5 hours). Mean attended 4.1 (SD=1.3) sessions (range 2-6) <sup>2</sup> 3x weekly 1-hour sessions (3 hours) <sup>3</sup>   |  |  |  |
| Comparator                               | Waitlist <sup>1,2</sup>   |  |  |  |
| Comparator                               | TAU: All participants were given the same milieu treatment, including weekly individual psychotherapy (provided primarily by Master's level therapists), weekly group psychotherapy, special education services, a behaviour modification point system, and, on an individual basis as needed, medication and/or psychoeducational parent/family counselling <sup>3</sup> |  |  |  |
| Intervention length (weeks)              | 8 <sup>1</sup><br>6 <sup>2</sup><br>3 <sup>3</sup>  |  |  |  |
|  |   |  |  |  |

- ADIS-C-Anxiety Disorders Interview Schedule-Child interview; ADHD-Attention Deficit Hyperactivity 1 2 3 4 5 Disorder; EMDR-Eye movement desensitisation and reprocessing; DSM-Diagnostic and Statistical
- Manual of Mental Disorders; ICD-International Classification of Disease; NR-Not reported; ODD-
- Oppositional defiant disorder; TAU-Treatment as usual
- <sup>1</sup>Ahmad 2007/2008; <sup>2</sup>de Roos 2017; <sup>3</sup>Soberman 2002
- 6 See appendix D for full evidence tables.

## 7 Quality assessment of clinical studies included in the evidence review

- 8 The clinical evidence profile for this review (EMDR for the treatment of PTSD in
- 9 children and young people) is presented in Table 17.

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# Table 17: Summary clinical evidence profile: EMDR versus waitlist or TAU for delayed treatment (>3 months)

| delaye  | ed treatmer                              | nt (>3 months)   |                                |  |  |
|---|--|--|--------------------------------|--|--|
|   | Illustrative comparative risks* (95% CI) |  |                                |  |  |
| Outcomes  | Assumed risk Waitlist or TAU             | Corresponding risk Eye movement desensitisation and reprocessing (EMDR)  | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| PTSD symptomatolo gy self-rated at endpoint CRTI/CRIES change score Follow-up: 3-6 weeks                                    | OI TAU                                   | The mean PTSD symptomatology self-rated at endpoint in the intervention groups was 0.9 standard deviations lower (2.64 lower to 0.85 higher)           | (33 / 601)                     | 82<br>(2 studies)                      | very<br>low <sup>1,2,3,4</sup>           |
| PTSD<br>symptomatolo<br>gy self-rated<br>at 2-month<br>follow-up<br>CRIES<br>change score<br>Follow-up:<br>mean 2<br>months |  | The mean PTSD symptomatology self-rated at 2-month follow-up in the intervention groups was 0.72 standard deviations lower (1.57 lower to 0.13 higher) |                                | 23<br>(1 study)                        | low <sup>1,5</sup>                       |
| PTSD<br>symptomatolo<br>gy clinician-<br>rated<br>PTSS-C<br>change score<br>Follow-up:<br>mean 8 weeks                      |  | The mean PTSD symptomatology clinician-rated in the intervention groups was 0.07 standard deviations higher (0.61 lower to 0.76 higher)                |                                | 33<br>(1 study)                        | very low <sup>1,3</sup>                  |
| Emotional and<br>behavioural<br>problems<br>SDQ-A<br>change score<br>Follow-up:<br>mean 6 weeks                             |  | The mean emotional and behavioural problems in the intervention groups was 1.52 standard deviations lower (2.14 to 0.91 lower)                         |                                | 61<br>(1 study)                        | very low <sup>1,4,6</sup>                |
| Quality of life<br>KIDSCREEN-<br>27: Global<br>HRQoL T-<br>scores;<br>change score<br>Follow-up:<br>mean 6 weeks            |  | The mean quality of life in the intervention groups was 0.81 standard deviations higher (0.24 to 1.38 higher)  |                                | 61<br>(1 study)                        | very low <sup>1,4,6</sup>                |

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| Illustrative co<br>(95% CI)  |                              | comparative risks*  |                                |  |  |
|--|------------------------------|---|--------------------------------|--|--|
| Outcomes   | Assumed risk Waitlist or TAU | Corresponding risk Eye movement desensitisation and reprocessing (EMDR) | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Better indicated by higher values  |                              |   |                                |  |  |
| Discontinuation Number of participants lost to follow-up for any reason Follow-up: 3-8 weeks | 122 per<br>1000              | 80 per 1000<br>(18 to 353)  | RR 0.65<br>(0.15 to<br>2.88)   | 123<br>(3 studies)                     | low <sup>3</sup>                         |

CI=confidence interval; CRIES= Children's Revised Impact of Event Scale; CRTI= Children's Response to Trauma Inventory; HRQoL=Health Related Quality of Life; KIDSCREEN-27= Health-related quality of life questionnaire for children, young people and their parents; PTSD=post-traumatic stress disorder; PTSS=Post-Traumatic Stress Symptom; RR=risk ratio; SDQ-A= Strength and Difficulties

12 See appendix F for full GRADE tables.

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## 14 Hypnotherapy: clinical evidence

### 15 Included studies

- One study of hypnotherapy for the treatment of PTSD in children and young people
- was identified for full-text review. This study could not be included.

## 18 Excluded studies

- 19 One study was reviewed at full text and excluded from this review because the
- 20 intervention was outside protocol (spiritual-hypnosis).
- 21 Studies not included in this review with reasons for their exclusions are provided in
- 22 Appendix K.

Questionnaires; SMD=standardised mean difference; TAU=treatment as usual;

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> Considerable heterogeneity (I2>80%)

<sup>&</sup>lt;sup>3</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>4</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>5</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>6</sup> OIS not met (N<400)

## 1 Psychodynamic therapies: clinical evidence

### 2 Included studies

- 3 Three studies of psychodynamic therapies for the treatment of PTSD in children and
- 4 young people were identified for full-text review. Of these 3 studies, 1 RCT (N=75)
- 5 was included in a single comparison.
- 6 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 7 symptoms, there were no included studies.
- 8 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 9 PTSD symptoms, 1 RCT (N=75) compared child-parent psychotherapy using play
- with parent training (case management and individual treatment for parent-only) 10
- (Lieberman 2005/2006/ Ghosh Ippen 2011 [one study reported across three papers]). 11

#### 12 Excluded studies

- 13 Two studies were reviewed at full text and excluded from this review because the
- 14 study was a non-RCT (no control group) or the comparison was outside protocol
- (within-class individual versus group). 15
- Studies not included in this review with reasons for their exclusions are provided in 16
- 17 Appendix K.

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## 18 Summary of clinical studies included in the evidence review

- 19 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 20 study evidence tables in Appendix D.
- 21 Table 18 provides a brief summary of the included study and evidence from this
- study is summarised in the clinical GRADE evidence profile below (Table 19). 22
- 23 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 24 study evidence tables in Appendix D.

## Table 18: Summary of included studies: Psychodynamic therapies for delayed treatment (>3 months)

| troutinone (* o montino)            |   |  |  |  |
|-------------------------------------|---|--|--|--|
| Comparison                          | Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) |  |  |  |
| Total no. of studies (N randomised) | 1 (75)  |  |  |  |
| Study ID                            | Lieberman 2005/2006/Ghosh Ippen 2011  |  |  |  |
| Country                             | US  |  |  |  |
| Diagnostic status                   | Clinically important PTSD symptoms (scoring above a threshold on validated scale)                                       |  |  |  |
| Mean months since onset of PTSD     | NR  |  |  |  |
| Mean age (range)                    | 4.1 (3-5)   |  |  |  |
| Sex (% female)                      | 52  |  |  |  |
| Ethnicity (% BME)                   | 91  |  |  |  |
| Coexisting conditions               | NR  |  |  |  |

|  | Child-parent psychotherapy using play versus parent training (case management and individual treatment for   |
|--|--|
| Comparison                               | parent-only)   |
| Mean months since traumatic event        | NR   |
| Type of traumatic event                  | Domestic violence: Children exposed to marital violence  |
| Single or multiple incident index trauma | Multiple   |
| Lifetime experience of trauma            | Multiple stressors, including exposure to community violence (47%), physical abuse (19%), sexual abuse (15%), or both (4%). During the study, 33% of the mothers reported new traumas that affected the dyad and 17% of the mothers reported either returning to their violent partners or entering a new violent relationship |
| Intervention details                     | Child-Parent Psychotherapy (CPP, following manual by Lieberman & Van Horn 2005) with case management plus treatment as usual in the community  |
| Intervention format                      | Individual/Family  |
| Intervention intensity                   | 50x weekly 1-hour sessions (50 hours). Mean sessions attended 32.09 (SD=15.20)   |
| Comparator                               | Parent training (case management and individual treatment for parent-only). 73% of mothers and 55% of children received individual treatment, and 45% received separate individual psychotherapy for both mother and child   |
| Intervention length (weeks)              | 50   |

- 1 NR-Not reported; CPP-Child-Parent Psychotherapy
- 2 See appendix D for full evidence tables.

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## 3 Quality assessment of clinical studies included in the evidence review

- The clinical evidence profile for this review (psychodynamic therapy for the treatment of PTSD in children and young people) is presented in Table 19.
  - Table 19: Summary clinical evidence profile: Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for delayed treatment (>3 months)

| 1.0011.10111  | or paront or   | ny, for actayed tre  | w (                                   | •,                                     |                                  |
|---|--|--|---------------------------------------|--|----------------------------------|
|   | Illustrative of (95% CI)   | omparative risks*  |                                       |  |                                  |
| Outcomes  | Assumed risk Parent training (case managem ent and individual treatment for parent-only) | Corresponding risk Child-parent psychotherapy using play         | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality of the evidenc e (GRADE) |
| PTSD<br>symptomatology<br>clinician-rated<br>DC 0-3; change |  | The mean PTSD symptomatology clinician-rated in the intervention |                                       | 65<br>(1 study)                        | very<br>low <sup>1,2,3</sup>     |

|   | Illustrative c<br>(95% CI)   | omparative risks*   |                                       |  |                                 |
|---|--|---|---------------------------------------|--|---------------------------------|
| Outcomes  | Assumed risk Parent training (case managem ent and individual treatment for parent-only) | Corresponding risk Child-parent psychotherapy using play  | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality of the evidenc e (GRADE |
| score<br>Follow-up: mean 50<br>weeks  |  | groups was<br>1.19 standard<br>deviations lower<br>(1.72 to 0.66<br>lower)  |                                       |  |                                 |
| Emotional and<br>behavioural<br>problems at<br>endpoint<br>CBCL total; change<br>score<br>Follow-up: mean 50<br>weeks           |  | The mean emotional and behavioural problems at endpoint in the intervention groups was 0.79 standard deviations lower (1.3 to 0.28 lower)           |                                       | 65<br>(1 study)                        | very<br>low <sup>1,3</sup>      |
| Emotional and<br>behavioural<br>problems at 6-<br>month follow-up<br>CBCL total; change<br>score<br>Follow-up: mean 6<br>months |  | The mean emotional and behavioural problems at 6-month follow-up in the intervention groups was 0.98 standard deviations lower (1.58 to 0.39 lower) |                                       | 50<br>(1 study)                        | very<br>low <sup>1,2,3</sup>    |
| Discontinuation Number of participants lost to follow-up for any reason Follow-up: mean 50 weeks                                | 121 per<br>1000  | 143 per 1000<br>(44 to 465)   | RR<br>1.18<br>(0.36<br>to<br>3.84)    | 75<br>(1 study)                        | low <sup>4</sup>                |

CBCL=Children's Behavioural Checklist; CI=confidence interval; DC=Diagnostic Criteria; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference 123456

#### 7 See appendix F for full GRADE tables.

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

## 1 Counselling: clinical evidence

### 2 Included studies

- 3 Five studies of counselling for the treatment of PTSD in children and young people
- 4 were identified for full-text review. Of these 5 studies, 2 RCTs (N=125) were included
- 5 in a single comparison for counselling.
- 6 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 7 symptoms, there were no included studies.
- 8 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 9 PTSD symptoms, 2 RCTs (N=125) compared supportive counselling with no
- treatment or waitlist (Chen 2014; Ertl 2011/Neuner 2007 [published paper and
- 11 protocol]).
- 12 Comparison with trauma-focused CBT are presented in the Trauma-focused CBT
- 13 section above.

### 14 Excluded studies

- 15 Three studies were reviewed at full text and excluded from this review because the
- 16 paper was a commentary or book section.
- 17 Studies not included in this review with reasons for their exclusions are provided in
- 18 Appendix K.

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## 19 Summary of clinical studies included in the evidence review

- 20 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 21 study evidence tables in Appendix D.
- Table 20 provides brief summaries of the included studies and evidence from these
- are summarised in the clinical GRADE evidence profile below (Table 21).
- See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 25 study evidence tables in Appendix D.

## Table 20: Summary of included studies: Counselling for delayed treatment (>3 months)

| Comparison                          | Supportive counselling versus no treatment or waitlist   |
|-------------------------------------|--|
| Total no. of studies (N randomised) | 2 (125)  |
| Study ID                            | Chen 2014 <sup>1</sup><br>Ertl 2011/Neuner 2007 <sup>2</sup>   |
| Country                             | China <sup>1</sup><br>Uganda <sup>2</sup>  |
| Diagnostic status                   | Clinically important PTSD symptoms (scoring above a threshold on validated scale) <sup>1</sup> PTSD diagnosis according to ICD/DSM criteria <sup>2</sup> |
| Mean months since onset of PTSD     | NR   |
| Mean age (range)                    | 14.5 (range NR) <sup>1</sup> 18.4 (12-25) <sup>2</sup>   |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Comparison                               | Supportive counselling versus no treatment or waitlist  |
|--|---|
| Sex (% female)                           | 68 <sup>1</sup>   |
| ,  | 55 <sup>2</sup>   |
| Ethnicity (% BME)                        | NR  |
| Coexisting conditions                    | NR  |
| Mean months since traumatic event        | 24 <sup>1</sup><br>80.5 <sup>2</sup>  |
|  |   |
| Type of traumatic event                  | Natural disaster: Adolescents who had lost at least one parent in the Sichuan, China, Earthquake <sup>1</sup>   |
|  | Child soldiers: The duration of abduction ranged from several hours to 7.42 years, with a median of 2.47 months. The likelihood of an event being indicated as the worst if present was highest for being forced to kill (55%), followed by witnessed killing (31%) and seeing someone being mutilated or seeing dead bodies (13%) <sup>2</sup> |
| Single or multiple incident index trauma | Single <sup>1</sup> Multiple <sup>2</sup>   |
| Lifetime experience of                   | NR¹   |
| trauma                                   | Other than abduction, the most common traumatic event types reported by 81 or more of the 85 participants were exposure to a war zone, witnessing someone being killed, witnessing abduction, witnessing physical assault, and assaults with weapons <sup>2</sup>   |
| Intervention details                     | General support provided on an individual basis adopting counselling techniques such as listening, reflection, and empathy <sup>1</sup>   |
|  | Needs-based intervention incorporating an academic catch-<br>up program for just over half of the intervention time and with<br>the rest of the time equally dedicated to psychoeducation,<br>conducting discussions on coping with symptoms, and<br>dealing with current problems <sup>2</sup>   |
| Intervention format                      | Individual  |
| Intervention intensity                   | 6x weekly sessions (length of session NR) <sup>1</sup> 8x thrice-weekly 90-120-min sessions (12-16 hours) <sup>2</sup>  |
| Comparator                               | No treatment <sup>1</sup> Waitlist <sup>2</sup>   |
| Intervention length (weeks)              | 6 <sup>1</sup> 3 <sup>2</sup>   |
| ND Not remarked                          |   |

- NR-Not reported.
- 1 <sup>1</sup>Chen 2014; <sup>2</sup>Ertl 2011/Neuner 2007
- 3 See appendix F for full evidence tables.

## 4 Quality assessment of clinical studies included in the evidence review

- The clinical evidence profile for this review (counselling for the treatment of PTSD in 5
- 6 children and young people) is presented in Table 21.

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# Table 21: Summary clinical evidence profile: Supportive counselling versus no treatment or waitlist for delayed treatment (>3 months)

| treatment or waitlist for delayed treatment (>3 months)  |                                       |   |                                |                           |                         |
|--|---------------------------------------|---|--------------------------------|---------------------------|-------------------------|
|  |                                       | comparative risks*  |                                |                           |                         |
|  | (95% CI) Assumed risk No treatment or | Corresponding risk Supportive   | Relativ<br>e<br>effect<br>(95% | No of<br>Participant<br>s | Quality of the evidence |
| Outcomes   | waitlist                              | counselling   | CI)                            | (studies)                 | (GRADE)                 |
| PTSD<br>symptomatology<br>self-rated at<br>endpoint<br>CRIES change<br>score<br>Follow-up: mean<br>6 weeks               |                                       | The mean PTSD symptomatology self-rated at endpoint in the intervention groups was 0.48 standard deviations lower (1.33 lower to 0.37 higher)               |                                | 22<br>(1 study)           | low <sup>1,2</sup>      |
| PTSD symptomatology self-rated at 3-month follow-up CRIES change score Follow-up: mean 3 months                          |                                       | The mean PTSD symptomatology self-rated at 3-month follow-up in the intervention groups was 0.42 standard deviations lower (1.27 lower to 0.43 higher)      |                                | 22<br>(1 study)           | low <sup>1,2</sup>      |
| PTSD<br>symptomatology<br>clinician-rated at<br>3-month follow-up<br>CAPS change<br>score<br>Follow-up: mean<br>3 months |                                       | The mean PTSD symptomatology clinician-rated at 3-month follow-up in the intervention groups was 0.43 standard deviations lower (0.98 lower to 0.12 higher) |                                | 52<br>(1 study)           | low <sup>2,3</sup>      |
| PTSD<br>symptomatology<br>clinician-rated at<br>6-month follow-up<br>CAPS change<br>score<br>Follow-up: mean<br>6 months |                                       | The mean PTSD symptomatology clinician-rated at 6-month follow-up in the intervention groups was 0.11 standard deviations lower (0.66 lower to 0.44 higher) |                                | 51<br>(1 study)           | low <sup>2,3</sup>      |
| PTSD<br>symptomatology<br>clinician-rated at<br>12-month follow-<br>up<br>CAPS change<br>score                           |                                       | The mean PTSD symptomatology clinician-rated at 12-month follow-up in the intervention groups was 0 standard  |                                | 51<br>(1 study)           | very low <sup>3,4</sup> |

|   | Illustrative (95% CI)        | comparative risks*  |                                 |                     |                              |
|---|------------------------------|---|---------------------------------|---------------------|------------------------------|
| O. 4  | Assumed risk No treatment or | Corresponding risk Supportive   | Relativ<br>e<br>effect<br>(95%  | No of Participant s | Quality of the evidence      |
| Outcomes Follow-up: mean 12 months  | waitlist                     | deviations higher<br>(0.55 lower to 0.55<br>higher)   | CI)                             | (studies)           | (GRADE)                      |
| Remission at 12-<br>month follow-up<br>Number of people<br>no longer meeting<br>diagnostic criteria<br>for PTSD<br>Follow-up: mean<br>12 months | 536 per<br>1000              | 466 per 1000<br>(273 to 788)  | RR<br>0.87<br>(0.51 to<br>1.47) | 56<br>(1 study)     | very low <sup>3,4</sup>      |
| Depression<br>symptoms at<br>endpoint<br>CES-D change<br>score<br>Follow-up: mean<br>6 weeks  |                              | The mean depression symptoms at endpoint in the intervention groups was 0.11 standard deviations higher (0.73 lower to 0.95 higher)         |                                 | 22<br>(1 study)     | very low <sup>1,4</sup>      |
| Depression<br>symptoms at 3-<br>month follow-up<br>CES-<br>D/MINI:Depressio<br>n change score<br>Follow-up: mean<br>3 months                    |                              | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.7 standard deviations lower (1.17 to 0.22 lower)         |                                 | 74<br>(2 studies)   | very<br>low <sup>1,3,5</sup> |
| Depression<br>symptoms at 6-<br>month follow-up<br>MINI:Depression<br>change score<br>Follow-up: mean<br>6 months                               |                              | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.47 standard deviations lower (1.03 lower to 0.09 higher) |                                 | 51<br>(1 study)     | very<br>low <sup>1,2,3</sup> |
| Depression<br>symptoms at 12-<br>month follow-up<br>MINI:Depression<br>change score<br>Follow-up: mean<br>12 months                             |                              | The mean depression symptoms at 12-month follow-up in the intervention groups was 0.34 standard deviations lower                            |                                 | 51<br>(1 study)     | very<br>low <sup>1,2,3</sup> |

|   | Illustrative<br>(95% CI)              | comparative risks*   |                                       |  |  |
|---|---------------------------------------|--|---------------------------------------|--|--|
| Outcomes  | Assumed risk No treatment or waitlist | Corresponding risk Supportive counselling (0.9 lower to 0.21   | Relativ<br>e<br>effect<br>(95%<br>CI) | No of<br>Participant<br>s<br>(studies) | Quality of<br>the<br>evidence<br>(GRADE) |
| Functional impairment at 3-month follow-up CAPS: Functional impairment; change score Follow-up: mean 3 months   |                                       | higher) The mean functional impairment at 3-month follow-up in the intervention groups was 0.91 standard deviations lower (1.49 to 0.34 lower) |                                       | 52<br>(1 study)                        | low <sup>3,5</sup>                       |
| Functional impairment at 6-month follow-up CAPS: Functional impairment; change score Follow-up: mean 6 months   |                                       | The mean functional impairment at 6-month follow-up in the intervention groups was 0.44 standard deviations lower (1 lower to 0.12 higher)     |                                       | 51<br>(1 study)                        | low <sup>2,3</sup>                       |
| Functional impairment at 12-month follow-up CAPS: Functional impairment; change score Follow-up: mean 12 months |                                       | The mean functional impairment at 12-month follow-up in the intervention groups was 0.27 standard deviations lower (0.82 lower to 0.28 higher) |                                       | 51<br>(1 study)                        | low <sup>2,3</sup>                       |
| Discontinuation<br>Number of<br>participants lost<br>to follow-up for<br>any reason<br>Follow-up: 3-6<br>weeks  | 0 per<br>1000                         | 0 per 1000<br>(0 to 0)   | RR<br>6.75<br>(0.86 to<br>52.7)       | 80<br>(2 studies)                      | moderate <sup>6</sup>                    |

CAPS= Clinician Administered PTSD Symptom; CES-D= Centre for Epidemiological Studies-

123456789 Depression; CI=confidence interval; CRIES= Children's Revised Impact of Event Scale; PTSD=posttraumatic stress disorder; RR=risk ratio; SMD=standardised mean difference

#### 10 See appendix F for full GRADE tables.

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>5</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>6</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

## 1 Combined somatic and cognitive therapies: clinical evidence

### 2 Included studies

- 3 one study of a combined somatic and cognitive therapy for the treatment of PTSD in
- 4 children and young people was identified for full-text review and this RCT (N=60) was
- 5 included in a single comparison.
- 6 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 7 symptoms, there were no included studies.
- 8 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 9 PTSD symptoms, 1 RCT (N=60) compared emotional freedom technique (EFT) with
- 10 no treatment (Al-Hadethe 2015).
- 11 Comparison with trauma-focused CBT are presented in the Trauma-focused CBT
- 12 section above.

#### 13 Excluded studies

22

23

No studies were reviewed at full text and excluded from this review.

## 15 Summary of clinical studies included in the evidence review

- 16 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 17 study evidence tables in Appendix D.
- Table 22 provides a brief summary of the included study and evidence from this
- 19 study is summarised in the clinical GRADE evidence profile below (Table 23).
- 20 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 21 study evidence tables in Appendix D.

## Table 22: Summary of included studies: Combined somatic and cognitive therapies for delayed treatment (>3 months)

|  | ayed treatment (20 months)                            |
|--|---|
| Comparison                               | Emotional freedom technique (EFT) versus no treatment |
| Total no. of studies (N randomised)      | 1 (60)  |
| Study ID                                 | Al-Hadethe 2015                                       |
| Country                                  | Iraq  |
| Diagnostic status                        | PTSD diagnosis according to ICD/DSM criteria          |
| Mean months since onset of PTSD          | NR  |
| Mean age (range)                         | Mean NR (16-19)                                       |
| Sex (% female)                           | 0   |
| Ethnicity (% BME)                        | NR  |
| Coexisting conditions                    | NR  |
| Mean months since traumatic event        | NR  |
| Type of traumatic event                  | Unclear (details NR)                                  |
| Single or multiple incident index trauma | Unclear   |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Comparison                    | Emotional freedom technique (EFT) versus no treatment   |
|-------------------------------|---|
| Lifetime experience of trauma | NR  |
| Intervention details          | Emotional Freedom Technique (EFT). Acupressure-based treatment involving the participant tapping on specific meridian points while talking through traumatic memories |
| Intervention format           | Individual  |
| Intervention intensity        | 4x biweekly 60-90 min sessions (4-6 hours)  |
| Comparator                    | No treatment  |
| Intervention length (weeks)   | 2   |
| Note. None                    |   |

- 1 EFT-Emotional Freedom Technique; PTSD-Post-traumatic stress disorder; NR-Not reported.
- 2 See appendix D for full evidence tables.

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## 3 Quality assessment of clinical studies included in the evidence review

- 4 The clinical evidence profile for this review (combined somatic and cognitive therapy
- for the treatment of PTSD in children and young people) is presented in Table 23.

Table 23: Summary clinical evidence profile: Combined somatic and cognitive therapies versus no treatment for delayed treatment (>3 months)

|  | Illustrative comparative risks* (95% CI) |   |                                | Ì                                      |  |
|--|--|---|--------------------------------|--|--|
| Outcomes   | Assumed risk No treatment                | Corresponding risk Combined somatic and cognitive therapies   | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatolo<br>gy self-rated<br>at endpoint<br>SPTSS<br>change score<br>Follow-up:<br>mean 2<br>weeks |  | The mean PTSD symptomatology self-rated at endpoint in the intervention groups was 1.85 standard deviations lower (2.6 to 1.1 lower)            |                                | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| PTSD symptomatolo gy self-rated at 3-month follow-up SPTSS change score Follow-up: mean 3 months               |  | The mean PTSD symptomatology self-rated at 3-month follow-up in the intervention groups was 1.96 standard deviations lower (2.72 to 1.19 lower) |                                | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| PTSD symptomatolo gy self-rated at 6-month follow-up SPTSS change score Follow-up:                             |  | The mean PTSD symptomatology self-rated at 6-month follow-up in the intervention groups was 1.3 standard  |                                | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |

|  | Illustrative<br>(95% CI)  | comparative risks*   |                                |   |  |
|--|---------------------------|--|--------------------------------|---|--|
| Outcomes   | Assumed risk No treatment | Corresponding risk Combined somatic and cognitive therapies  | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies)  | Quality<br>of the<br>evidence<br>(GRADE) |
| mean 6<br>months   |                           | deviations lower<br>(1.99 to 0.61 lower)   | (1111)                         | (====================================== | ,  |
| PTSD symptomatolo gy self-rated at 12-month follow-up SPTSS change score Follow-up: mean 12 months           |                           | The mean PTSD symptomatology self-rated at 12-month follow-up in the intervention groups was 1.85 standard deviations lower (2.6 to 1.1 lower) |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>endpoint<br>HADS-A<br>change score<br>Follow-up:<br>mean 2<br>weeks                |                           | The mean anxiety symptoms at endpoint in the intervention groups was 0.95 standard deviations lower (1.61 to 0.3 lower)                        |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |
| Anxiety symptoms at 3-month follow-up HADS-A change score Follow-up: mean 3 months                           |                           | The mean anxiety symptoms at 3-month follow-up in the intervention groups was 0.89 standard deviations lower (1.54 to 0.24 lower)              |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>6-month<br>follow-up<br>HADS-A<br>change score<br>Follow-up:<br>mean 6<br>months   |                           | The mean anxiety symptoms at 6-month follow-up in the intervention groups was 1.15 standard deviations lower (1.82 to 0.47 lower)              |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |
| Anxiety<br>symptoms at<br>12-month<br>follow-up<br>HADS-A<br>change score<br>Follow-up:<br>mean 12<br>months |                           | The mean anxiety symptoms at 12-month follow-up in the intervention groups was 1.19 standard deviations lower (1.86 to 0.51 lower)             |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |
| Depression<br>symptoms at<br>endpoint<br>HADS-D  |                           | The mean depression symptoms at endpoint in the  |                                | 40<br>(1 study)                         | very<br>low <sup>1,2</sup>               |

|   | Illustrative<br>(95% CI)  | comparative risks*  |  |  |  |
|---|---------------------------|---|--|--|--|
| Outcomes  | Assumed risk No treatment | Corresponding risk Combined somatic and cognitive therapies   | Relative<br>effect<br>(95% CI)         | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| change score<br>Follow-up:<br>mean 2<br>weeks   |                           | intervention groups<br>was<br>0.94 standard<br>deviations lower<br>(1.59 to 0.28 lower)   | (, , , , , , , , , , , , , , , , , , , | <b></b>                                | (  |
| Depression<br>symptoms at<br>3-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:<br>mean 3<br>months   |                           | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.75 standard deviations lower (1.4 to 0.11 lower)   |  | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Depression<br>symptoms at<br>6-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:<br>mean 6<br>months   |                           | The mean depression symptoms at 6-month follow-up in the intervention groups was 0.85 standard deviations lower (1.5 to 0.2 lower)    |  | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Depression<br>symptoms at<br>12-month<br>follow-up<br>HADS-D<br>change score<br>Follow-up:<br>mean 12<br>months |                           | The mean depression symptoms at 12-month follow-up in the intervention groups was 1.38 standard deviations lower (2.07 to 0.68 lower) |  | 40<br>(1 study)                        | very<br>low <sup>1,2</sup>               |
| Discontinuation Number of participants lost to follow- up for any reason Follow-up: mean 2 weeks                | -                         | -   | Not<br>estimable                       | 40<br>(1 study)                        | low <sup>1,3</sup>                       |

CI=confidence interval; HADS-A/D= Hospital Anxiety and Depression Scale-Anxiety/Depression; 123456 PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; SPTSS=

Screen for Post-Traumatic Stress Symptoms; TAU=treatment as usual <sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>2</sup> OIS not met (N<400)

<sup>3</sup> OIS not met (events<300)

7 See appendix F for full GRADE tables.

## 1 Parent training/family interventions: clinical evidence

### 2 Included studies

- 3 Four studies of parent training or family interventions for the treatment of PTSD in
- 4 children and young people were identified for full-text review. Of these 4 studies, 3
- 5 RCTs (N=286) were included in 3 comparisons for parent training/family
- 6 interventions.
- 7 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 8 symptoms, there were no included studies.
- 9 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 10 PTSD symptoms, 1 RCT (N=100) compared parent training (CBT with parent-only)
- 11 with TAU (Deblinger 1996/1999 [one study reported across 2 papers), 1 RCT (N=36)
- 12 compared parent training in addition to trauma-focused CBT for the child with
- trauma-focused CBT for the child-only (King 2000), and 1 RCT (N=150) compared
- family therapy with waitlist (Kazak 2004).
- 15 Comparison with trauma-focused CBT are presented in the Trauma-focused CBT
- 16 section above.

#### 17 Excluded studies

- 18 One study was reviewed at full text and excluded from this review because efficacy
- 19 or safety data could not be extracted.
- 20 Studies not included in this review with reasons for their exclusions are provided in
- 21 Appendix K.

## 22 Summary of clinical studies included in the evidence review

- 23 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 24 study evidence tables in Appendix D.
- Table 24 provides brief summaries of the included studies and evidence from these
- are summarised in the clinical GRADE evidence profiles below (Table 25, Table 26
- 27 and Table 27).

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- See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 29 study evidence tables in Appendix D.

## Table 24: Summary of included studies: Parent training/family interventions for delayed treatment (>3 months)

| Comparison                          | Parent training (CBT with parent-only) versus TAU         | Parent training + TF-<br>CBT (for child) versus<br>TF-CBT (for child | Family therapy versus waitlist                            |
|-------------------------------------|---|--|---|
| Total no. of studies (N randomised) | 1 (100)   | 1 (36)   | 1 (150)   |
| Study ID                            | Deblinger 1996/1999                                       | King 2000  | Kazak 2004  |
| Country                             | US  | Australia  | US  |
| Diagnostic status                   | Clinically important<br>PTSD symptoms<br>(scoring above a | Clinically important PTSD symptoms (scoring above a                  | Clinically important<br>PTSD symptoms<br>(scoring above a |

|   | Doront training (CDT   | Doront training L.T.   | Eamily thorony   |
|---|--|--|--|
|   | Parent training (CBT with parent-only)   | Parent training + TF-<br>CBT (for child) versus  | Family therapy versus waitlist   |
| Comparison                                    | versus TAU   | TF-CBT (for child threshold on validated   | throphold on validated   |
|   | threshold on validated scale)  | scale)   | threshold on validated scale)  |
| Mean<br>months<br>since onset<br>of PTSD      | NR   | NR   | NR   |
| Mean age (range)                              | 9.8 (7-13)   | 11.4 (5-17)  | 14.6 (10-19)   |
| Sex (% female)                                | 83   | 69   | 52   |
| Ethnicity (% BME)                             | 28   | NR   | 12   |
| Coexisting conditions                         | 29% major depression;<br>30% oppositional<br>defiant disorder; 20%<br>ADHD; 11% separation<br>anxiety; 6% conduct<br>disorder; 5% specific<br>phobia; 1% OCD   | For 69% who met DSM-IV criteria for full PTSD (N=25): 16% with full PTSD had no other Axis I diagnoses, 36% had one comorbid diagnosis, 40% had two comorbid diagnoses, and 8% had three comorbid diagnoses. The comorbid diagnoses included dysthymia (28%), oppositional defiant disorder (28%), separation anxiety disorder (24%), generalized anxiety disorder (20%), conduct disorder (12%), major depression (8%), attentiondeficit/hyperactivity disorder (8%), and specific phobia (8%). | NR   |
| Mean<br>months<br>since<br>traumatic<br>event | Mean NR (for 66% the last sexually abusive incident occurred in the 6 months prior to initial assessment, 16% 6 months to 2 years before initial assessment, and 18% 2 more years before the evaluation) | 54.5   | 63.6 (SD=35.0) since completion of cancer treatment. The median age at diagnosis was 7.80 years (range=2.76 months to 16.36 years)     |
| Type of traumatic event                       | Childhood sexual abuse: Contact sexual abuse. 18% experienced 1 sexually abusive incident, 47% 2-10 episodes, 22%  | Childhood sexual abuse: In the majority of cases, the offenders were male adults known to the child such as the biological father,   | Diagnosis of life-<br>threatening condition:<br>Diagnoses included<br>leukaemia (25%), solid<br>tumours (22%),<br>lymphoma (21%), bone |

| Comparison  | Parent training (CBT with parent-only) versus TAU   | Parent training + TF-<br>CBT (for child) versus<br>TF-CBT (for child  | Family therapy versus waitlist  |
|---|---|---|---|
|   | 11-50 episodes, and<br>13% >50 abusive<br>incidents   | stepfather, family<br>friend, neighbour, or<br>teacher. Nearly all of<br>the children had<br>experienced multiple<br>episodes of sexual<br>abuse involving<br>penetration offenses<br>and other forms of<br>sexual abuse                  | tumours (8%), and other (24%)   |
| Single or<br>multiple<br>incident<br>index trauma | Multiple  | Multiple  | Single  |
| Lifetime<br>experience<br>of trauma               | NR  | Mean number of abusive episodes: 7.6 (SD=3.8; range 1-33)   | NR  |
| Intervention<br>details                           | Parents taught to respond therapeutically to child behaviour and needs using graded exposure, modelling, education and coping. Caregiver seen alone for full session, child invited in for evaluation purposes only | Parent training (modelled on Cohen & Mannarino 1996 and Deblinger & Heflin 1996) in child behaviour management skills and parent—child communication skills. The child received trauma-focused CBT, based on protocol of Deblinger (1996) | Surviving Cancer<br>Competently<br>Intervention Program<br>(SCCIP; following<br>manual by Kazak<br>1999)  |
| Intervention format                               | Individual  | Individual/Family   | Group   |
| Intervention intensity                            | 12x weekly 45-min<br>sessions (9 hours)   | 20x weekly 50-min<br>sessions (16.7 hours)<br>for child + 20x weekly<br>50-min sessions (16.7<br>hours) for parent (33.3<br>hours in total)   | 4-sessions in 1-day (5 hours of direct therapeutic contact and an additional 2 hours of informal contact during breaks). All families completed all four sessions |
| Comparator  | TAU: Parents and children were given information about symptom patterns and encouraged to access therapy, and child protection workers or the victim witness coordinator were asked to assist with referrals        | Trauma-focused CBT<br>for child, based on<br>protocol of Deblinger<br>(1996)  | Waitlist  |
| Intervention<br>length<br>(weeks)                 | 12  | 20  | 0.1   |

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- ADHD-Attention Deficit Hyperactivity Disorder; CBT-Cognitive Behaviour Therapy; DSM-Diagnostic and
- 1 2 3 Statistical Manual of Mental Disorders; NR-Not reported; SCCIP-Surviving Cancer Competently
- Interventions Program; TAU-Treatment as usual.
- 4 See appendix D for full evidence tables.

## 5 Quality assessment of clinical studies included in the evidence review

- The clinical evidence profiles for this review (parent training/family interventions for 6
- 7 the treatment of PTSD in children and young people) are presented in Table 25,
- Table 26 and Table 27. 8

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## Table 25: Summary clinical evidence profile: Parent training (CBT with parentonly) versus TAU for delayed treatment (>3 months)

| Oiliy)  |                 | or delayed treatmen   | it (> 3 iii0i                      | 11113)                             |  |
|---|-----------------|---|------------------------------------|------------------------------------|--|
|   | Illustrative co | omparative risks*   |                                    |                                    |  |
| Outcomes  | Assumed risk    | Corresponding risk Parent training (CBT with parent- only)  | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD symptomatolo gy clinician-rated at endpoint K-SADS-E: PTSD; change score Follow-up: mean 12 weeks          |                 | The mean PTSD symptomatology clinician-rated at endpoint in the intervention groups was 0.59 standard deviations lower (1.29 lower to 0.11 higher)          |                                    | 34<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| PTSD symptomatolo gy clinician-rated at 3-month follow-up K-SADS-E: PTSD; change score Follow-up: mean 3 months |                 | The mean PTSD symptomatology clinician-rated at 3-month follow-up in the intervention groups was 0.63 standard deviations lower (1.33 lower to 0.07 higher) |                                    | 34<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| PTSD symptomatolo gy clinician-rated at 6-month follow-up K-SADS-E: PTSD; change score Follow-up: mean 6 months |                 | The mean PTSD symptomatology clinician-rated at 6-month follow-up in the intervention groups was 0.58 standard deviations lower (1.28 lower to 0.12 higher) |                                    | 34<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| PTSD symptomatolo   |                 | The mean PTSD symptomatology  |                                    | 34<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |

|  | Illustrative co  | omparative risks*   |                                    |                                    |  |
|--|------------------|---|------------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk TAU | Corresponding risk Parent training (CBT with parent- only)  | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| gy clinician- rated at 12- month follow- up K-SADS-E: PTSD; change score Follow-up: mean 12 months   | TAU              | clinician-rated at 12-month follow-up in the intervention groups was 0.42 standard deviations lower (1.11 lower to 0.27 higher)   | Cij                                | (studies)                          | (GRADE)                                  |
| PTSD symptomatolo gy clinician-rated at 2-year follow-up K-SADS-E: PTSD; change score Follow-up: mean 2 years  |                  | The mean PTSD symptomatology clinician-rated at 2-year follow-up in the intervention groups was 0.89 standard deviations lower (1.6 to 0.17 lower)                      |                                    | 34<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Emotional and<br>behavioural<br>problems-<br>Externalizing<br>at endpoint<br>CBCL:<br>Externalizing;<br>change score<br>Follow-up:<br>mean 12<br>weeks |                  | The mean emotional and behavioural problems-externalizing at endpoint in the intervention groups was 0.63 standard deviations lower (1.38 lower to 0.12 higher)         |                                    | 30<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Emotional and behavioural problems- Externalizing at 3-month follow-up CBCL: Externalizing; change score Follow-up: mean 3 months                      |                  | The mean emotional and behavioural problems-externalizing at 3-month follow-up in the intervention groups was 0.23 standard deviations lower (0.96 lower to 0.5 higher) |                                    | 30<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |
| Emotional and behavioural problems- Externalizing at 6-month follow-up CBCL: Externalizing;  |                  | The mean emotional and behavioural problems-externalizing at 6-month follow-up in the intervention groups was 0.18 standard   |                                    | 30<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |

|   | Illustrative co | omparative risks*  |                                    |                                    |  |
|---|-----------------|--|------------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk    | Corresponding risk Parent training (CBT with parent- only)   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| change score Follow-up: mean 6 months   | IAU             | deviations lower<br>(0.91 lower to 0.55<br>higher)   | Oi)                                | (studies)                          | (GRADE)                                  |
| Emotional and behavioural problems- Externalizing at 12-month follow-up CBCL: Externalizing; change score Follow-up: mean 12 months |                 | The mean emotional and behavioural problems-externalizing at 12-month follow-up in the intervention groups was 0.07 standard deviations lower (0.8 lower to 0.66 higher) |                                    | 30<br>(1 study)                    | very<br>low <sup>1,3,5</sup>             |
| Emotional and behavioural problems- Externalizing at 2-year follow-up CBCL: Externalizing; change score Follow-up: mean 2 years     |                 | The mean emotional and behavioural problems-externalizing at 2-year follow-up in the intervention groups was 0.92 standard deviations lower (1.69 to 0.15 lower)         |                                    | 30<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression<br>symptoms at<br>endpoint<br>CDI change<br>score<br>Follow-up:<br>mean 12<br>weeks                                      |                 | The mean depression symptoms at endpoint in the intervention groups was 0.86 standard deviations lower (1.56 to 0.15 lower)  |                                    | 35<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression<br>symptoms at<br>3-month<br>follow-up<br>CDI change<br>score<br>Follow-up:<br>mean 3<br>months                          |                 | The mean depression symptoms at 3-month follow-up in the intervention groups was 0.45 standard deviations lower (1.13 lower to 0.23 higher)                              |                                    | 35<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Depression<br>symptoms at<br>6-month<br>follow-up<br>CDI change   |                 | The mean depression symptoms at 6-month follow-up in the intervention  |                                    | 35<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |

|  | Illustrative comparative risks* (95% CI) |   |                                    |                                    |  |
|--|--|---|------------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk                             | Corresponding risk<br>Parent training<br>(CBT with parent-<br>only)   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| score<br>Follow-up:<br>mean 6<br>months  |  | groups was<br>0.32 standard<br>deviations lower<br>(1 lower to 0.35<br>higher)  |                                    |                                    |  |
| Depression<br>symptoms at<br>12-month<br>follow-up<br>CDI change<br>score<br>Follow-up:<br>mean 12<br>months |  | The mean depression symptoms at 12-month follow-up in the intervention groups was 0.5 standard deviations lower (1.18 lower to 0.18 higher) |                                    | 35<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Depression<br>symptoms at<br>2-year follow-<br>up<br>CDI change<br>score<br>Follow-up:<br>mean 2 years       |  | The mean depression symptoms at 2-year follow-up in the intervention groups was 0.86 standard deviations lower (1.56 to 0.15 lower)         |                                    | 35<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |

CBCL=Children's Behavioural Checklist; CBT=cognitive behavioural therapy; CDI=Children's Depression Inventory; CI=confidence interval; K-SADS=Kiddele Schedulae for Affective Disorder and Schizophrenia; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; TAU=treatment as usual

<sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>4</sup> OIS not met (N<400)

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11 12 <sup>5</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

Table 26: Summary clinical evidence profile: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child)-only for delayed treatment (>3 months)

|  | Illustrative cor<br>(95% CI)                 | nparative risks*  |                                |                                    |  |
|--|--|---|--------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk Trauma-focused CBT (child only) | Correspondin g risk Trauma- focused CBT + parent training                               | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatolo<br>gy clinician-<br>rated at<br>endpoint<br>ADIS-C: |  | The mean PTSD symptomatology clinician-rated at endpoint in the intervention groups was |                                | 24<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |

|   | Illustrative con                             | nparative risks*   |                                |                                    |  |
|---|--|--|--------------------------------|------------------------------------|--|
| Outcomes  | Assumed risk Trauma-focused CBT (child only) | Correspondin<br>g risk<br>Trauma-<br>focused CBT<br>+ parent<br>training   | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD;<br>change score<br>Follow-up:<br>mean 20<br>weeks   |  | 0.36 standard<br>deviations lower<br>(1.16 lower to 0.4<br>higher)   | 5                              |                                    |  |
| PTSD symptomatolo gy clinician-rated at 3-month follow-up ADIS-C: PTSD; change score Follow-up: mean 3 months |  | The mean PTSD symptomatology clinician-rated at 3-month follow-up in the intervention groups was 0.48 standard deviations lower (1.29 lower to 0.3 higher) | n                              | 24<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Anxiety<br>symptoms at<br>endpoint<br>RCMAS;<br>change score<br>Follow-up:<br>mean 20<br>weeks                |  | The mean anxiety symptoms at endpoint in the intervention groups was 0.14 standard deviations higher (0.66 lower to 0.9 higher)                            |                                | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Anxiety symptoms at 3-month follow-up RCMAS; change score Follow-up: mean 3 months                            |  | The mean anxiety symptoms at 3-month follow-up is the intervention groups was 0.03 standard deviations higher (0.77 lower to 0.8 higher)                   | n                              | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression<br>symptoms at<br>endpoint<br>CDI; change<br>score<br>Follow-up:<br>mean 20<br>weeks               |  | The mean depression symptoms at endpoint in the intervention groups was 0.29 standard deviations lower (1.09 lower to 0.5 higher)                          | 2                              | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Depression symptoms at 3-month follow-up  |  | The mean depression symptoms at 3-month follow-up i  | n                              | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |

|  | Illustrative con                             | nparative risks*  |                                |                                    |  |
|--|--|---|--------------------------------|------------------------------------|--|
| Outcomes CDI; change score   | Assumed risk Trauma-focused CBT (child only) | Correspondin g risk Trauma- focused CBT + parent training the intervention groups was   | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Follow-up:<br>mean 3<br>months   |  | 0.07 standard<br>deviations lower<br>(0.87 lower to 0.7<br>higher)  | 3                              |                                    |  |
| Emotional and<br>behavioural<br>problems-<br>Internalizing<br>at endpoint<br>CBCL:<br>Internalizing;<br>change score<br>Follow-up:<br>mean 20<br>weeks             |  | The mean emotional and behavioural problems-internalizing at endpoint in the intervention groups was 0.29 standard deviations lower (1.1 lower to 0.51 higher)          |                                | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Emotional and<br>behavioural<br>problems-<br>Internalizing<br>at 3-month<br>follow-up<br>CBCL:<br>Internalizing;<br>change score<br>Follow-up:<br>mean 3<br>months |  | The mean emotional and behavioural problems-internalizing at 3-month follow-up in the intervention groups was 0.15 standard deviations lower (0.95 lower to 0.6 higher) | n                              | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Emotional and<br>behavioural<br>problems-<br>Externalizing<br>at endpoint<br>CBCL:<br>Externalizing;<br>change score<br>Follow-up:<br>mean 20<br>weeks             |  | The mean emotional and behavioural problems-externalizing at endpoint in the intervention groups was 0.79 standard deviations lower (1.63 lower to 0.0 higher)          | 4                              | 24<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Emotional and<br>behavioural<br>problems-<br>Externalizing<br>at 3-month<br>follow-up<br>CBCL:   |  | The mean emotional and behavioural problems-externalizing at 3 month follow-up i the intervention   |                                | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |

|  | Illustrative cor<br>(95% CI)                 | nparative risks*  |                                |                                    |  |
|--|--|---|--------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk Trauma-focused CBT (child only) | Correspondin<br>g risk<br>Trauma-<br>focused CBT<br>+ parent<br>training  | Relative<br>effect<br>(95% CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| Externalizing;<br>change score<br>Follow-up:<br>mean 3<br>months   |  | groups was<br>0.14 standard<br>deviations lower<br>(0.94 lower to 0.6<br>higher)  | 37                             |                                    |  |
| Global functioning at endpoint GAF; change score Follow-up: mean 20 weeks Better indicated by higher values          |  | The mean global functioning at endpoint in the intervention groups was 0.3 standard deviations higher (0.5 lower to 1.11 higher)          |                                | 24<br>(1 study)                    | very<br>low <sup>1,3,4</sup>             |
| Global functioning at 3-month follow-up GAF; change score Follow-up: mean 3 months Better indicated by higher values |  | The mean global functioning at 3-month follow-up i the intervention groups was 0.66 standard deviations higher (0.16 lower to 1.4 higher) | n                              | 24<br>(1 study)                    | very<br>low <sup>1,2,3</sup>             |
| Discontinuation  Number of participants lost to follow-up for any reason Follow-up: mean 20 weeks                    | 250 per 1000                                 | 250 per 1000<br>(62 to 1000)  | RR 1<br>(0.25 to<br>4)         | 24<br>(1 study)                    | low <sup>4</sup>                         |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS=; RR=risk ratio; SMD=standardised mean difference

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<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

 $<sup>^4</sup>$  95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

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Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## Table 27: Summary clinical evidence profile: Family therapy versus waitlist for delayed treatment (>3 months)

| delayed treatment (>3 months)   |  |  |                                 |                         |                       |  |
|---|--|--|---------------------------------|-------------------------|-----------------------|--|
|   | Illustrative comparative risks* (95% CI) |  | Relativ                         | No of                   | Quality of            |  |
| Outcomes  | Assume<br>d risk<br>Waitlist             | Corresponding risk Family therapy  | e effect<br>(95%<br>CI)         | Participant s (studies) | the evidence (GRADE)  |  |
| PTSD<br>symptomatology<br>UCLA PTSD-RI;<br>change score<br>Follow-up: mean<br>0.1 weeks           |  | The mean PTSD symptomatology in the intervention groups was 0.37 standard deviations lower (0.7 to 0.05 lower)       |                                 | 149<br>(1 study)        | low <sup>1,2</sup>    |  |
| Anxiety symptoms<br>RCMAS; T-scores<br>change score<br>Follow-up: mean<br>0.1 weeks               |  | The mean anxiety symptoms in the intervention groups was 0.09 standard deviations higher (0.24 lower to 0.41 higher) |                                 | 149<br>(1 study)        | low <sup>1,2</sup>    |  |
| Discontinuation Number of participants lost to follow-up for any reason Follow-up: mean 0.1 weeks | 68 per<br>1000                           | 382 per 1000<br>(156 to 932)   | RR<br>5.65<br>(2.31 to<br>13.8) | 150<br>(1 study)        | moderate <sup>3</sup> |  |

- CI=confidence interval; PTSD=post-traumatic stress disorder; RCMAS=Revised Children Manifest
  - Anxiety Scale; RR=risk ratio; SMD=standardised mean difference; UCLA PTSD-RI=UCLA PTSD-
- Reaction Index:
- 345678 <sup>1</sup> Risk of bias is high or unclear across multiple domains
- <sup>2</sup> OIS not met (N<400)
- <sup>3</sup> OIS not met (events<300)
- 9 See appendix F for full GRADE tables.

## Play therapy: clinical evidence

## 11 Included studies

- 12 Two studies of play therapy for the treatment of PTSD in children and young people
- were identified for full-text review, and both RCTs (N=162) were included. There 13
- were 2 comparisons for play therapy. 14
- 15 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 16 symptoms, there were no included studies.
- 17 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- PTSD symptoms, 1 RCT (N=131) compared play therapy with TAU (Deeba & Rapee 18
- 2015), and 1 RCT (N=31) compared play therapy with trauma-focused CBT 19
- 20 (Schottelkorb 2012).

### 21 Excluded studies

No studies were reviewed at full text and excluded from this review. 22

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Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## 1 Summary of clinical studies included in the evidence review

- See also the study selection flow chart in Appendix C, forest plots in Appendix E and study evidence tables in Appendix D.
- 4 Table 28 provides brief summaries of the included studies and evidence from these
- are summarised in the clinical GRADE evidence profiles below (Table 29 and Table 30).
- See also the study selection flow chart in Appendix C, forest plots in Appendix E and study evidence tables in Appendix D.

# Table 28: Summary of included studies: Play therapy for delayed treatment (>3 months)

| months)   |   |   |  |  |  |  |
|---|---|---|--|--|--|--|
| Comparison  | Play therapy versus TAU   | Play therapy versus trauma-<br>focused CBT  |  |  |  |  |
| Total no. of studies (N randomised)               | 1 (131)   | 1 (31)  |  |  |  |  |
| Study ID  | Deeba 2015  | Schottelkorb 2012   |  |  |  |  |
| Country   | Bangladesh  | US  |  |  |  |  |
| Diagnostic status                                 | Clinically important PTSD symptoms (scoring above a threshold on validated scale)   | Clinically important PTSD symptoms (scoring above a threshold on validated scale)         |  |  |  |  |
| Mean months since onset of PTSD                   | NR  | NR  |  |  |  |  |
| Mean age<br>(range)                               | 7.2 (5-9)   | 9.2 (6-13)  |  |  |  |  |
| Sex (% female)                                    | 37  | 45  |  |  |  |  |
| Ethnicity (%<br>BME)                              | NR  | 67  |  |  |  |  |
| Coexisting conditions                             | NR  | NR  |  |  |  |  |
| Mean months since traumatic event                 | NR (time in shelter home: 40.3 months)  | NR  |  |  |  |  |
| Type of traumatic event                           | Mixed: Children living in a shelter home. Most of the children (90%) had lost one or both parents following natural disasters or accidents or due to domestic violence and witnessed direct or indirect violence against a parent (mostly towards the mother) | Witnessing war as a civilian:<br>Childhood refugee trauma (no<br>further detail reported) |  |  |  |  |
| Single or<br>multiple<br>incident index<br>trauma | Multiple  | Multiple  |  |  |  |  |
| Lifetime experience of trauma                     | NR  | NR  |  |  |  |  |

| Comparison                  | Play therapy versus TAU   | Play therapy versus trauma-<br>focused CBT   |
|-----------------------------|---|--|
| Intervention details        | Enhanced Huggy Puppy<br>Intervention, following similar<br>protocol to Sadeh (2008) | Child-centred play therapy (CCPT; following the manual by Ray 2011)  |
| Intervention format         | Individual/Family   | Individual/Family  |
| Intervention intensity      | 2x sessions (length of sessions NR)   | 24x biweekly 30-min sessions (12 hours) + 6x 15-min parent consultation sessions (1.5 hours) (13.5 hours in total). Mean 17 sessions completed + 3 sessions with parents |
| Comparator                  | TAU (no further detail reported)  | Cohen TF-CBT/Cognitive processing therapy (based on the manual by Cohen 2006), with child only and parent only portions and conjoint parent—child sessions               |
| Intervention length (weeks) | 3   | 12   |

- BME-Black and minority ethnic; CBT-Cognitive Behaviour Therapy; CCPT-Child centred play therapy;
- 1 NR-Not reported; PTSD-Post-traumatic stress disorder; TAU-Treatment as usual.
- 3 See appendix D for full evidence tables.

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## 4 Quality assessment of clinical studies included in the evidence review

- 5 The clinical evidence profiles for this review (play therapy for the treatment of PTSD
- in children and young people) are presented in Table 29 and Table 30. 6

Table 29: Summary clinical evidence profile: Play therapy versus TAU for delayed treatment (>3 months)

|   | Illustrative comparative risks* (95% CI) |   | Relativ                 |                              | Quality                       |
|---|--|---|-------------------------|------------------------------|-------------------------------|
| Outcomes  | Assumed risk TAU                         | Corresponding risk Play therapy   | e effect<br>(95%<br>CI) | No of Participants (studies) | of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatology<br>self-rated<br>CRIES change<br>score<br>Follow-up:<br>mean 3 weeks |  | The mean PTSD symptomatology self-rated in the intervention groups was 1.07 standard deviations lower (1.44 to 0.7 lower) |                         | 129<br>(1 study)             | very<br>low <sup>1,2,3</sup>  |
| Anxiety<br>symptoms<br>SCASp; change<br>score<br>Follow-up:<br>mean 3 weeks                 |  | The mean anxiety symptoms in the intervention groups was 1.87 standard deviations lower (2.29 to 1.45 lower)              |                         | 129<br>(1 study)             | very<br>low <sup>1,2,3</sup>  |
| Depression<br>symptoms<br>SMFQp; change   |  | The mean depression symptoms in the   |                         | 129<br>(1 study)             | very<br>low <sup>1,2,3</sup>  |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

|   | Illustrative comparative risks* (95% CI) |   | Relativ                      |                              | Quality                       |
|---|--|---|------------------------------|------------------------------|-------------------------------|
| Outcomes  | Assumed risk TAU                         | Corresponding risk Play therapy   | e effect<br>(95%<br>CI)      | No of Participants (studies) | of the<br>evidence<br>(GRADE) |
| score<br>Follow-up:<br>mean 3 weeks   |  | intervention groups<br>was<br>1.34 standard<br>deviations lower<br>(1.73 to 0.96 lower) |                              |                              |                               |
| Discontinuation Number of participants lost to follow-up for any reason Follow-up: mean 3 weeks | 32 per<br>1000                           | 6 per 1000<br>(0 to 119)  | RR 0.18<br>(0.01 to<br>3.68) | 131<br>(1 study)             | very<br>low <sup>1,4</sup>    |

- CI-confidence interval; CRIES; PTSD-post-traumatic stress disorder; RR-risk ratio; SCAS-Spence Children's Anxiety Scale; SMD-standardised mean difference; SMFQ-Short Mood and Feeling
- 1234567 Questionnaires; TAU-treatment as usual
- <sup>1</sup> Risk of bias is high or unclear across multiple domains
- <sup>2</sup> OIS not met (N<400)

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- <sup>3</sup> Data is not reported/cannot be extracted for all outcomes
  - <sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

Table 30: Summary clinical evidence profile: Play therapy versus traumafocused CBT for delayed treatment (>3 months)

|  | Illustrative comparative risks* (95% CI) |   |                                    |                                    |  |
|--|--|---|------------------------------------|------------------------------------|--|
| Outcomes   | Assumed risk Trauma-focused CBT          | Corresponding risk Play therapy   | Relativ<br>e effect<br>(95%<br>CI) | No of<br>Participants<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatolog<br>y self-rated<br>UCLA PTSD-RI;<br>change score<br>Follow-up:<br>mean 12 weeks              |  | The mean PTSD symptomatology self-rated in the intervention groups was 0.11 standard deviations lower (0.88 lower to 0.66 higher) |                                    | 26<br>(1 study)                    | very<br>low <sup>1,2</sup>               |
| Discontinuation<br>Number of<br>participants lost<br>to follow-up for<br>any reason<br>Follow-up:<br>mean 12 weeks | 294 per<br>1000                          | 32 per 1000<br>(3 to 535)   | RR 0.11<br>(0.01 to<br>1.82)       | 31<br>(1 study)                    | low <sup>2</sup>                         |

- 10 CI=confidence interval; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean
- 11 difference; TAU=treatment as usual; UCLA PTSD-RI=UCLA PTSD-Reaction Index.
- 12 13 <sup>1</sup> Risk of bias is high or unclear across multiple domains
- <sup>2</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm
- 14 See appendix F for full GRADE tables.

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## 1 Self-help (without support): clinical evidence

#### 2 Included studies

- 3 One study of self-help (without support) for the treatment of PTSD in children and
- 4 young people was identified for full-text review. This study could not be included.

#### 5 Excluded studies

- 6 One study was reviewed at full text and excluded from this review because the paper
- 7 was a systematic review with no new useable data and any meta-analysis results not
- 8 appropriate to extract.
- 9 Studies not included in this review with reasons for their exclusions are provided in
- 10 Appendix K.

### 11 Economic evidence

#### 12 Included studies

- 13 Four studies assessing the cost effectiveness of psychological interventions for the
- treatment of children and young people with PTSD were identified (Gospodarevskaya
- and Segal, 2012; McCrone 2005; Mihalopoulos 2015; Shearer 2018). The search
- strategy for economic studies is provided in Appendix B.

## 17 Excluded studies

- 18 No economic studies of psychological interventions for the treatment of children and
- 19 young people with PTSD were reviewed at full text and excluded.

## 20 Summary of studies included in the economic evidence review

- 21 Gospodarevskaya and Segal (2012) developed a decision-analytic economic model
- 22 to assess the cost effectiveness of trauma-focused CBT, alone or in combination with
- 23 SSRIs, compared with non-directive supportive counselling and no treatment for
- 24 children with PTSD in Australia. The modelled population comprised 10-year-old
- 25 children who met all or most of the PTSD diagnostic criteria, including at least one
- symptom of avoidance or re-experiencing; some of the children had comorbid
- 27 depression. Both psychological interventions comprised 12 weekly sessions lasting
- 28 45 minutes each. The analysis adopted a mental health system perspective. Costs
- included staff's time (psychologist, psychiatrist, GP, social worker), SSRI acquisition
- 30 costs and parental group or psychoeducational sessions over 12 months; beyond 12
- 31 months, only antidepressant and GP costs were considered for children with
- 32 recurrent depression. Efficacy data were taken from meta-analyses of RCTs and
- 33 further adjustments via indirect comparisons. Resource use data were based on trial
- information; national unit costs were used. The measure of outcome was the QALY,
- 35 estimated using utility scores elicited from the Australian population using the
- 36 Assessment of Quality of Life (AQoL-4D) instrument. The time horizon of the analysis
- was 31 years. During this period, it was assumed that, following successful
- treatment, no relapses of PTSD due to the original traumatic event occurred; in
- 39 contrast, the model allowed the recurrence of depressive episodes, which were
- 40 treated with SSRIs.
- The most effective intervention was the combination of trauma-focused CBT with
- 42 SSRIs. Counselling was found to be less effective and more costly than trauma-

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Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

focused CBT alone. The incremental cost effectiveness ratio (ICER) of combined 2 therapy versus trauma-focused CBT alone was Aus\$2,901/QALY in 2011 prices 3 (£1,442/QALY in 2016 prices). The ICER of trauma-focused CBT versus no 4 treatment was Aus\$1,650/QALY in 2011 prices (£820/QALY in 2016 prices). Using 5 the NICE lower cost effectiveness threshold of £20,000/QALY, combination therapy 6 of trauma-focused CBT and SSRIs was the most cost-effective intervention. Results 7 were sensitive to variation in clinical effectiveness, as expected. The study is partially 8 applicable to the NICE decision-making context as it was conducted in Australia and 9 the method of QALY estimation is not consistent with NICE recommendations (NICE 10 recommends use of EQ-5D and the UK utility value tariff). The study is characterised 11 by potentially serious limitations, including its narrow perspective and some 12 modelling assumptions (the model assumed no relapses of PTSD over 31 years).

Mihalopoulos and colleagues (2015) undertook a model-based cost-utility analysis to compare trauma-focused CBT (consisting of 8-10 individual sessions delivered by a psychologist) with non-evidence-based treatment as usual, comprising consultation with healthcare professionals for children and young people with PTSD in Australia. The eligible study population comprised prevalent cases (12-month prevalence) of children with PTSD in Australia in 2012, who were currently seeking care, had consulted any health professional for a mental health problem during the previous 12 months but had not been receiving evidence-based care. The perspective of the analysis was that of the health sector (government and service user out-of-pocket expenses). Only intervention costs were included (psychologist's, psychiatrist's or GP's time). Efficacy data were taken from meta-analysis of trial data. Resource use data were based on trial and epidemiological data and expert opinion; national unit costs were used. The measure of outcome was the QALY, estimated using utility scores elicited from the Australian population using the Assessment of Quality of Life (AQoL-4D) instrument. The Disability-Adjusted Life Year (DALY) was also used. The time horizon of the analysis was 5 years; a 3% annual discount rate was used. However, only benefits were measured for a period of 5 years (assuming that benefits are retained over this period); costs were measured over the duration of treatment (i.e. up to 8-10 weeks).

32 Trauma-focused CBT was found to be more costly and more effective than treatment 33 as usual, with an ICER of Aus\$8,900/QALY in 2012 prices (£3,954/QALY in 2016 34 prices). The probability of trauma-focused CBT being cost-effective was 1 at a 35 willingness to pay of \$50,000/QALY (£22,214/QALY). Results were most sensitive to 36 PTSD prevalence, effectiveness of trauma-focused CBT, adherence and eligibility for 37 CBT. The study is partially applicable to the NICE decision-making context as it was 38 conducted in Australia and the method of QALY estimation is not consistent with 39 NICE recommendations. The study is characterised by potentially serious limitations, 40 including the short time used for measuring costs (until end of treatment) and the fact 41 that only intervention costs (therapist's time) were considered.

42 McCrone and colleagues (2005) estimated the costs of short-term individual 43 psychodynamic psychotherapy (up to 30 sessions) and psychoeducational group therapy (up to 18 sessions) assessed in a RCT (Trowel 2002); the trial participants 44 45 were sexually abused girls 6-14 years old, with symptoms of emotional or behavioural disturbance, 73% of whom had PTSD. Both interventions included 46 47 carers' support. The analysis was conducted from the perspective of providers of 48 mental health services to children and support to parents. Only intervention costs 49 were considered, comprising therapists' time, including introductory meeting, initial 50 assessment, therapy, carers' support, supervision of therapists and carers' workers, 51 and follow up assessments for up to 2 years from treatment initiation. Efficacy and resource use data were based on the RCT (N=75; at 1-year follow up: n=58; at 2-52

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- 1 year follow up: n=54); national unit costs were used. The outcome measures of the
- 2 RCT were the global impairment of functioning measured using the K-GAS and the
- 3 Orvaschel's PTSD scale.
- 4 The authors conducted the study as a cost-minimisation analysis, as they reported
- 5 that results between the two interventions were similar. However, psychodynamic
- 6 psychotherapy showed greater improvements in manifestations of PTSD compared
- 7 with psychoeducational group therapy, with an effect size ranging from 0.60 to 0.79.
- 8 Psychodynamic psychotherapy was found to be significantly more expensive than
- 9 psychoeducational group therapy, with a cost difference of £2,051 per person treated
- in 2016 prices. The study is partially applicable to the NICE decision-making context
- as it used a narrow perspective and did not use QALYs as the outcome measure.
- 12 The study is characterised by potentially serious limitations, including its narrow
- perspective and the lack of synthesis of costs and outcomes.
- 14 Shearer and colleagues (2018) conducted an economic evaluation of trauma-focused
- 15 cognitive therapy (which belongs to the class of TF-CBT) versus waitlist for children
- and adolescents with PTSD in the UK. The analysis was based on the results of a
- 17 11-week RCT (Meiser-Stedman 2010/2017, N=29), which were extrapolated over a
- period of 3 year using decision-analytic economic modelling. Trauma-focused
- 19 cognitive therapy comprised 10 weekly individual sessions delivered by a trained
- 20 clinical psychologist. The analysis adopted a NHS and Personal Social Services
- 21 (PSS) perspective. Costs included intervention costs (psychologist's time), hospital
- 22 care (inpatient, outpatient, emergency department, ambulance), community health
- and social care staff time (GP, GP nurse, district nurse, paediatrician, clinical
- 24 psychologist, CAMHS worker, counsellor, educational psychologist), advice service,
- social and other services, and medication. Efficacy and cost data were based on the
- 26 RCT; national unit costs were used. The measure of outcome was the QALY,
- 27 estimated using the Strengths and Difficulties Questionnaire (SDQ), which was then
- 28 mapped onto the preference-based Child Health Utility index 9 dimensions (CHU-
- 9D) that has been valued by a sample of Australian population of young people.
- 30 During the 3-year time horizon of the model it was assumed that no relapses of
- 31 PTSD due to the original traumatic event occurred.
- Trauma-focused cognitive therapy was found to be more costly and more effective
- than waitlist, with an ICER of £2,205/QALY in 2014 prices (£2,254/QALY in 2016
- prices). The probability of trauma-focused cognitive therapy being cost-effective was
- 35 0.60-0.69 at a willingness-to-pay (WTP) of £20,000-£30,000/QALY, respectively. In a
- 36 completer case analysis, the ICER increased only slightly, at £2,806/QALY in 2014
- 37 prices (£2,869 in 2016 prices), with the probability of trauma-focused cognitive
- therapy being cost-effective reaching 0.69-0.75 at a WTP of £20,000-£30,000/QALY,
- 39 respectively. When psychologist training costs were included in the analysis, the
- 40 ICER of trauma-focused cognitive therapy versus waitlist rose up to £16,187/QALY
- 41 (£16,549 in 2016 prices); the probability of trauma-focused cognitive therapy being
- 42 cost-effective fell at 0.51-0.62 at a WTP of £20,000-£30,000/QALY, respectively. The
- 43 study is partially applicable to the NICE decision-making context, because, although
- it was conducted in the UK and adopted a NHS/PSS perspective, the method of
- 45 QALY estimation is not consistent with NICE recommendations. The study is
- 46 characterised by potentially serious limitations, mainly that costs and efficacy data
- were derived from a small RCT (N=29) with a short duration (11 weeks).
- The references of included studies and the economic evidence tables are provided in
- 49 Appendix H. The economic evidence profiles are shown in Appendix I.

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

### 1 Economic model

- 2 A decision-analytic model was developed to assess the relative cost effectiveness of
- 3 psychological interventions for the treatment of PTSD in children and young people.
- 4 The objective of economic modelling, the methodology adopted, the results and the
- 5 conclusions from this economic analysis are described in detail in Appendix J. This
- 6 section provides a summary of the methods employed and the results of the
- 7 economic analysis.

## 8 Overview of economic modelling methods

- 9 A hybrid decision-analytic model consisting of a decision-tree followed by a two-state
- 10 Markov model was constructed to evaluate the relative cost effectiveness of a range
- of psychological interventions for the treatment of children and young people with
- 12 clinically important symptoms of PTSD in a community setting. The time horizon of
- the analysis was 3 years, consisting of the 6 months of the decision tree and another
- 14 2.5 years (10 x 3-month cycles) in the Markov component of the economic model.
- 15 The range of interventions assessed in the economic analysis was determined by the
- availability of relevant clinical data included in the guideline systematic review of
- interventions for the treatment of children and young people with clinically important
- symptoms of PTSD. Network meta-analysis (NMA) was employed for synthesis of
- 19 the available efficacy data. The guideline economic analysis assessed psychological
- 20 interventions that were connected to the network of evidence and were thus possible
- 21 to include in the NMA. The NMA and the economic analysis considered separately
- 22 interventions that belonged to the trauma-focused cognitive behavioural therapy (TF-
- 23 CBT) class, as individual interventions had different intervention costs and,
- 24 potentially, different efficacy. Based on the advice of the committee, only effective
- interventions that had been tested on at least 40 people across the RCTs included in
- the NMAs assessing efficacy at treatment endpoint were considered in the economic
- analysis, as this was deemed as the minimum evidence that would be adequate to
- support a practice recommendation. The only exception was cognitive therapy, which
- 29 had been tested on only 25 children; this was included in the economic analysis as
- 30 the committee was interested in the relative clinical and cost effectiveness across all
- 31 interventions belonging in the TF-CBT class, and sufficient evidence on TF-CBT
- 32 class, which could be extrapolated to cognitive therapy, was available for other
- interventions within the class.
- 34 Based on the available evidence, the following interventions were considered in the
- 35 economic analysis of psychological interventions for the treatment of children and
- 36 young people with clinically important symptoms of PTSD:
- Supportive counselling
- Group CBT (TF-CBT)
- Cohen TF-CBT / Cognitive processing therapy [Cohen/CPT] (TF-CBT)
- Cognitive therapy (TF-CBT)
- Narrative exposure (TF-CBT)
- Exposure /prolonged exposure (TF-CBT)
- Eve Movement Desensitisation Reprocessing [EMDR]
- Family therapy
- 45 Play therapy
- Parent training
- No treatment, reflected in waitlist or no treatment arms of RCTs included in the
   guideline systematic review and NMA.

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Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

According to the model structure, hypothetical cohorts of children and young people with clinically important symptoms of PTSD were initiated on each of the treatment options assessed, including no treatment. Following a course of treatment, children and young people in each cohort either remitted (that is, they did not meet criteria for a PTSD diagnosis) or did not remit. In the next 3 months of follow-up after end of treatment, those who remitted ('no PTSD') could remain in remission or relapse to a PTSD state. Conversely, those who did not remit, could remain in the PTSD state or could remit (and move to a 'no PTSD' state). After that point, children and young people in each cohort were entered into the Markov component of the economic model, in either the 'PTSD' or the 'no PTSD' health states, depending on their state at the end of the decision-tree. In each cycle of the Markov model, they could remain in the same health state or move between the two states of 'PTSD' and 'no PTSD'.

13 Efficacy data were derived from the guideline systematic review and NMAs. The 14 baseline risk of remission was determined based on a review of published evidence; 15 the risk of relapse was based on the committee's expert opinion. The measure of 16 outcome of the economic analysis was the number of QALYs gained. Utility data 17 were selected after a systematic review of the literature. The perspective of the 18 analysis was that of health and personal social care services. Resource use was 19 based on published literature. National UK unit costs were used. The cost year was 20 2017. Model input parameters were synthesised in a probabilistic analysis. This 21 approach allowed more comprehensive consideration of the uncertainty 22 characterising the input parameters and captured the non-linearity characterising the 23 economic model structure.

The main analysis utilised efficacy data at treatment endpoint from a NMA of continuous data (changes in PTSD symptom scores), transformed to log-odds ratios of remission; a secondary analysis used efficacy data at treatment endpoint from a NMA of dichotomous remission data. Four scenarios were explored in probabilistic analysis:

- Scenario A (base-case analysis) utilised base-case utility data and assumed no beneficial effect of interventions beyond treatment endpoint
- Scenario B utilised base-case utility data and efficacy data at 3 months posttreatment from a NMA of continuous data (changes in PTSD symptom scores) between baseline and 1-4 month follow-up, transformed to log-odds ratios of remission; the secondary analysis used the odds ratio of group CBT versus waitlist at 1-4 month follow-up to estimate the relative effect of all interventions versus no treatment at 3-6 months.
- Scenario C utilised alternative utility data and assumed no beneficial effect of interventions beyond treatment endpoint
- Scenario D utilised alternative utility data and efficacy data at 3 months posttreatment from a NMA of continuous data (changes in PTSD symptom scores) between baseline and 1-4 month follow-up, transformed to log-odds ratios of remission; the secondary analysis used the odds ratio of group CBT versus waitlist at 1-4 month follow-up to estimate the relative effect of all interventions versus no treatment at 3-6 months.
- One-way deterministic sensitivity analysis was employed to explore the impact of a change in the annual risk of relapse.
- 47 Results have been expressed in the form of Incremental Cost Effectiveness Ratios
- 48 (ICERs) following the principles of incremental analysis. Net Monetary Benefits
- 49 (NMBs) have also been estimated. Incremental mean costs and effects (QALYs) of
- each intervention versus no treatment have been presented in the form of cost

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- 1 effectiveness planes. Results of probabilistic analysis have been summarised in the
- form of cost effectiveness acceptability curves (CEACs), which express the
- 3 probability of each intervention being cost effective at various cost effectiveness
- 4 thresholds. Cost effectiveness acceptability frontiers (CEAFs) have also been plotted;
- 5 these show the treatment option with the highest mean NMB over different cost
- 6 effectiveness thresholds, and the probability that the option with the highest NMB is
- 7 the most cost-effective among those assessed.

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## Overview of economic modelling results and conclusions

- 9 In the base-case analysis (which utilised base-case utility data at treatment endpoint
- and assumed no treatment effect beyond treatment endpoint), the order of
- interventions from the most to the least cost-effective for the treatment of PTSD in
- 12 children and young people was: cognitive therapy (TF-CBT), narrative exposure (TF-
- 13 CBT), play therapy, prolonged exposure (TF-CBT), Cohen/CPT (TF-CBT), EMDR,
- parent training, group CBT (TF-CBT), family therapy, supportive counselling and no
- treatment. The probability of cognitive therapy being the most cost-effective
- treatment option was 0.78. In the secondary analysis that used dichotomous
- 17 remission data, the order of interventions from the most to least cost-effective was:
- narrative exposure (TF-CBT), cognitive therapy (TF-CBT), prolonged exposure (TF-
- 19 CBT), Cohen/CPT (TF-CBT), no treatment and supportive counselling. The
- 20 probability of narrative exposure being the most cost-effective option was 0.62.
- 21 When a beneficial effect of up to 3 months post-treatment was assumed, the relative
- 22 cost effectiveness of group CBT and Cohen/CPT (both TF-CBT) improved and the
- 23 cost effectiveness of play therapy was reduced. The order of interventions became
- cognitive therapy (TF-CBT), Cohen/CPT (TF-CBT), group CBT (TF-CBT), narrative
- exposure (TF-CBT), parent training, prolonged exposure (TF-CBT), play therapy,
- 26 EMDR, supportive counselling, family therapy, no treatment. The probability of
- 27 cognitive therapy being the most cost-effective treatment option was 0.67. In the
- secondary analysis, the cost effectiveness of all interventions improved. Narrative
- 29 exposure remained the most cost-effective intervention with a 0.81 probability.
- followed by cognitive therapy and then prolonged exposure.
- 31 When narrower utility benefits for remission and no beneficial effect beyond
- 32 treatment endpoint were assumed, less costly interventions, such as EMDR and
- 33 group CBT, were favoured so that their relative cost effectiveness improved. The top-
- 34 3 most cost-effective interventions remained the same with those of the base-case
- analysis and the order of interventions by cost effectiveness was as follows: cognitive
- therapy (TF-CBT), narrative exposure (TF-CBT), play therapy, group CBT (TF-CBT),
- 37 EMDR, prolonged exposure (TF-CBT), parent training, Cohen/CPT (TF-CBT), family
- therapy, no treatment, supportive counselling. The probability of cognitive therapy
- 39 being the most cost-effective treatment option was 0.59. In secondary analysis, only
- 40 narrative exposure and cognitive therapy were more cost-effective than no treatment;
- 41 the probability of narrative exposure being the most cost-effective option was 0.71.
- When narrower utility benefits for remission and a beneficial effect up to 3 months
- post-treatment were assumed, the order of interventions from most to least cost-
- 44 effective became: cognitive therapy (TF-CBT), group CBT (TF-CBT), Cohen/CPT
- 45 (TF-CBT), narrative exposure (TF-CBT), parent training, EMDR, play therapy,
- 46 prolonged exposure (TF-CBT), family therapy, no treatment and supportive
- 47 counselling. The probability of cognitive therapy being the most cost-effective
- 48 intervention was only 0.31. In secondary analysis, the order of interventions by cost
- 49 effectiveness was: narrative exposure, Cohen/CPT, prolonged exposure, cognitive

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- therapy, supportive counselling, and no treatment. The probability of narrative
- 2 exposure being the most cost-effective option was 0.79.
- 3 Results of the economic analysis were overall robust to the changes in the risk of
- 4 relapse tested in deterministic sensitivity analysis.
- 5 Overall, individual forms of TF-CBT and, to a lesser degree, play therapy appear to
- 6 be cost-effective in the treatment of children and young people with PTSD. Family
- 7 therapy and supportive counselling do not appear to be cost-effective relative to other
- 8 interventions and, under some scenarios, supportive counselling is less cost-effective
- 9 than no treatment. In-between, there is another group of interventions (EMDR, group
- 10 CBT and parent training) with modest relative cost effectiveness, which is affected by
- the alternative scenarios tested. The secondary analysis confirmed the cost
- 12 effectiveness of individual forms of TF-CBT versus supportive counselling and no
- treatment, although the limited evidence did not allow further comparisons to be
- 14 made.
- 15 The results of the NMAs of 1-4 month follow-up PTSD change score data and of the
- dichotomous remission data showed considerable uncertainty due to the small size
- of the included studies and the small total number of studies. Thus, results based on
- these data should be interpreted with caution. Nevertheless, the base-case economic
- analysis did not utilise the outputs of any of these NMAs. The NMA that informed the
- 20 base-case economic analysis was based on more robust data and was characterised
- 21 by moderate heterogeneity and no evidence of inconsistency.

## 22 Resource impact

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- 23 The committee made a number of recommendations based on this review. Three of
- them were weaker ('consider') recommendations. Unlike for stronger ('offer')
- 25 recommendations that interventions should be adopted, it is not possible to make a
- 26 judgement about the potential resource impact to the NHS, as uptake of interventions
- 27 is difficult to predict. Overall, recommendations based on this review are not
- 28 expected to have a substantial impact on resources.
- 29 The committee's considerations that contributed to the resource impact assessment
- 30 are included under the 'Cost effectiveness and resource use' in 'The committee's
- 31 discussion of the evidence' section.

### 32 Clinical evidence statements

## 33 Trauma-focused CBT for early treatment (1-3 months)

- Low to moderate quality single-RCT (N=30-31) evidence suggests non-significant differences between trauma-focused CBT and meditation, for PTSD
- symptomatology or the number of participants with a PTSD diagnosis at endpoint
- or 6-month follow-up or on the rate of discontinuation, for children and young
- people with PTSD who were exposed to trauma 1-3 months ago.

## 39 Trauma-focused CBT for delayed treatment (>3 months)

- Very low to low quality evidence from 7-13 RCTs (N=409-872) suggests a large
   and statistically significant benefit of trauma-focused CBT relative to waitlist, TAU
- or no treatment on improving PTSD symptomatology (self-rated and clinician-
- rated) in children and young people with PTSD who were exposed to trauma more
- than 3 months ago. Very low to low quality evidence from 1-5 RCTs (N=39-301)
- suggests these benefits are maintained up to 12-month follow-up, but not at 18-
- 46 month or 2-year follow-ups (longest follow-up available for self-rated and clinician-

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- rated respectively). Moderate to very low quality evidence from 2-5 RCTs (N=90-277) also suggests clinically important and statistically significant benefits of trauma-focused CBT on the rate of remission and response, and benefits on remission are maintained at 1-3 month follow-up but not 12-18 month follow-up (longest follow-up available, no follow-up data available for response). In addition, low to very low quality evidence from 1-13 RCTs (N=39-834) suggests moderateto-large and statistically significant benefits on anxiety and depression symptoms. and emotional and behavioural problems, that are maintained up to 12-18 month follow-up (but not up to 2-year follow-up for depression). Very low quality evidence from 2 RCTs (N=219) also suggests a small but statistically significant benefit of trauma-focused CBT on improving quality of life. Very low quality evidence from 1-2 RCT analyses (N=53-220) suggests a clinically important benefit that just misses statistical significance of trauma-focused CBT for improving functional impairment at endpoint, and large and statistically significant benefit at 3-month and 12-month follow-up (effect at 6-month follow-up non-significant). Very low to low quality evidence from 1-4 RCTs (N=24-321) suggests large and statistically significant benefits on global functioning that are maintained up to 3-month follow-up but not at 18-month follow-up (longest follow-up available). Finally, moderate quality evidence from 18 RCTs (N=1255) suggests a trend for higher discontinuation with trauma-focused CBT, however, this effect is not statistically significant.
- Sub-analyses of the trauma-focused CBT relative to waitlist, TAU or no treatment comparison suggests some differential relative efficacy primarily for the clinicianrated PTSD symptomatology outcome (with the exception of format where subgroup differences are observed on self-rated PTSD symptomatology). Subanalysis by specific intervention suggested largest benefits observed for cognitive therapy and narrative exposure therapy, however benefits are clinically important and statistically significant across specific interventions. Sub-analysis by multiplicity of trauma also suggests differential efficacy with relatively larger benefits observed for children and young people who have been exposed to single incident index trauma, however, benefits for children and young people who have been exposed to multiple incident index trauma are also clinically important and statistically significant. Sub-analysis by format of intervention also suggests some differential efficacy with relatively larger effects observed for individual (child-only) and caregiver and child relative to group interventions, however, benefits are clinically important and statistically significant across formats. Sub-analysis by diagnostic status at baseline also suggests some differential efficacy with relatively larger effects observed for those with a diagnosis of PTSD at baseline relative to those with clinically important PTSD symptoms (who may not necessarily have a diagnosis). However, again effects are clinically important and statistically significant for both subgroups. Sub-analysis by trauma type also suggests some differences in effect sizes with relatively larger benefits observed for children and young people who have experienced motor vehicle accidents or witnessed war as a civilian, although clinically important and statistically significant benefits are observed across all trauma types. Finally, sub-analysis by age range (studies that include children aged under 7 years relative to studies where all children are over 7) suggests non-significant subgroup differences.
- Very low to moderate quality evidence from 2-5 RCTs (N=109-325) suggests
  moderate and statistically significant benefits of trauma-focused CBT relative to
  supportive counselling on improving PTSD symptomatology (self-rated and
  clinician-rated) in children and young people with PTSD who were exposed to
  trauma more than 3 months ago, and these benefits are maintained at longest
  follow-up available (12-17 months for self-rated and 12-months for clinician-rated).
   Very low to moderate quality evidence from 1-4 RCTs (N=61-208) also suggests
  clinically important and statistically significant benefits on the rate of remission and

- response and improving global functioning, that are maintained at 12-month follow-up (longest follow-up available). Very low quality single-RCT (N=48-82) analyses suggests moderate-to-large and statistically significant delayed benefits of trauma-focused CBT on improving dissociative symptoms and functional impairment (significant at 6- and 12-month follow-up, and just 12-month follow-up respectively). Conversely, very low to low quality evidence from 2-7 RCTs (N=70-552) suggests small but statistically significant benefits of trauma-focused CBT on anxiety and depression symptoms, but these benefits are short-term (nonsignificant at 6- and 12-month follow-up, and 3-, 6- and 12-17 month follow-up for anxiety and depression symptoms respectively). Very low quality evidence from 2-3 RCTs (N=224-385) suggests non-significant differences between traumafocused CBT and supportive counselling for internalising and externalising and behaviour problems at endpoint, 6- or 12-month follow-up. Moderate quality evidence from 8 RCTs (N=678) suggests a trend for a higher discontinuation rate associated with counselling relative to trauma-focused CBT, however this effect is not statistically significant. Sub-analyses by multiplicity of trauma, specific intervention, intervention format, age range, diagnostic status at baseline and trauma type suggest non-significant subgroup differences.
- Very low quality single-RCT (N=24) evidence suggests large and statistically significant benefits of a combined trauma-focused CBT and parent training intervention relative to waitlist on improving clinician-rated PTSD symptomatology and global functioning at endpoint and 3-month follow-up for children and young people with PTSD who were exposed to trauma more than 3 months ago. Evidence from this RCT suggests a large and statistically significant delayed benefit of a combined trauma-focused CBT and parent training intervention on emotional and behavioural problems (non-significant at endpoint and significant at 3-month follow-up). Whereas, evidence from this same RCT suggests non-significant effects on anxiety or depression symptoms. Low quality evidence from this RCT suggests a trend for a higher rate of discontinuation associated with a combined trauma-focused CBT and parent training intervention relative to waitlist, however this effect is not statistically significant.
- Low quality single-RCT (N=65-159) evidence suggests moderate and statistically significant benefits of a combined trauma-focused CBT and psychoeducational group intervention relative to psychoeducational group-only on self-rated PTSD symptomatology, the rate of response and depression symptoms at endpoint and 4-month follow-up for children and young people with PTSD who were exposed to trauma more than 3 months ago. Moderate quality evidence from this same RCT (N=159) suggests a trend for a higher rate of discontinuation associated with psychoeducational group-only, however this effect is not statistically significant.

### 40 Non-trauma-focused CBT for delayed treatment (>3 months)

Very low quality single-RCT (N=33) evidence suggests a large and statistically significant benefit of non-trauma-focused CBT in addition to TAU relative to TAU-only for improving substance use disorder symptoms in adolescents and young people with PTSD who were exposed to trauma more than 3 months ago.
 However, evidence from this same RCT suggests this benefit is not maintained at 3-month follow-up, effects on depression symptoms and discontinuation are non-significant, and no PTSD outcomes are available.

# 48 Eye movement desensitisation and reprocessing (EMDR) for delayed treatment (>3 months)

 Very low to low quality evidence from 1-2 RCTs (N=23-82) suggests clinically important but not statistically significant benefits of EMDR relative to waitlist or 

- TAU on self-rated PTSD symptomatology at endpoint and 2-month follow-up, and large and statistically significant benefits of EMDR on improving emotional and behavioural problems and quality of life, for children and young people with PTSD who were exposed to trauma more than 3 months ago. However, very low quality evidence from another RCT (N=33) suggests non-significant effects of EMDR on clinician-rated PTSD symptomatology. Low quality evidence from 3 RCTs (N=123) suggests a trend for a lower rate of discontinuation associated with EMDR relative to waitlist or TAU, however, this effect is not statistically significant.
  - Very low to low quality evidence from 1-2 RCTs (N=48-133) suggests a non-significant difference between trauma-focused CBT and EMDR on self-rated PTSD symptomology and quality of life (at endpoint, 3- and 12-month follow-up), clinician-rated PTSD symptomatology (at endpoint, no follow-up available) and discontinuation, for children and young people with PTSD who were exposed to trauma more than 3 months ago. Very low quality single-RCT (N=85) evidence suggests a small-to-moderate and statistically significant benefit of EMDR relative to trauma-focused CBT on emotional and behavioural problems at endpoint and 3- and 12-month follow-up.

## 18 Psychodynamic therapies for delayed treatment (>3 months)

Very low quality single-RCT (N=50-65) evidence suggests a large and statistically significant benefit of child-parent psychotherapy using play relative to parent training (case management and individual treatment for parent-only) on improving clinician-rated PTSD symptomatology at endpoint (no follow-up available) and emotional and behavioural problems at endpoint and 6-month follow-up, for children and young people with PTSD who were exposed to trauma more than 3 months ago. Low quality evidence from this same RCT (N=75) suggests non-significant effects on discontinuation.

## 27 Counselling for delayed treatment (>3 months)

• Very low to low quality evidence from single-RCT (N=22-56) analyses suggests non-significant effects of supportive counselling relative to no treatment or waitlist on self-rated PTSD symptomatology (at endpoint or 3-month follow-up), clinician-rated PTSD symptomatology (at 3-, 6- or 12-month follow-up) or remission (at 12-month follow-up), for children and young people with PTSD who were exposed to trauma more than 3 months ago. Very low quality evidence from 2 RCTs (N=74) suggests a moderate and statistically significant benefit of supportive counselling on depression symptoms at 3-month follow-up, however, effects at endpoint (K=1; N=22), 6-month and 12-month follow-up (K=1; N=51) are non-significant. Low quality evidence from 1 of these RCTs (N=52) also suggests a large and statistically significant benefit of supportive counselling on improving functional impairment at 3-month follow-up, however, again effects are not maintained at longer-term follow-up (6-month and 12-month). Moderate quality evidence from 2 RCTs (N=80) suggests a trend for a higher rate of discontinuation associated with supportive counselling, however this effect is not statistically significant.

## 43 Combined somatic and cognitive therapies for delayed treatment (>3 months)

 Very low quality single-RCT (N=40) evidence suggests large and statistically significant benefits of emotional freedom technique (EFT) relative to no treatment on improving self-rated PTSD symptomatology, anxiety and depression symptoms and benefits are maintained up to 12-month follow-up, for children and young people with PTSD who were exposed to trauma more than 3 months ago. No participants discontinued from this study.  • Very low quality single-RCT (N=39) evidence suggests large and statistically significant benefits of emotional freedom technique (EFT) relative to traumafocused CBT on self-rated PTSD symptomatology that is maintained up to 12-month follow-up, for children and young people with PTSD who were exposed to trauma more than 3 months ago. Evidence from the same RCT suggests large and statistically significant short-term benefits of EFT on improving anxiety symptoms (significant at endpoint and 3-month follow-up and non-significant at 6-and 12-month follow-up). Large or moderate and statistically significant benefits of EFT relative to trauma-focused CBT are also shown on depression symptoms at endpoint and 12-month follow-up (but non-significant at 3- and 6-month follow-up). Evidence from this RCT suggests a trend for a higher discontinuation rate associated with trauma-focused CBT relative to EFT, however this effect is not statistically significant.

## 14 Parent training/family interventions for delayed treatment (>3 months)

- Low quality single-RCT (N=149) evidence suggests a small but statistically significant benefit of family therapy relative to waitlist on improving PTSD symptomatology for children and young people with PTSD who were exposed to trauma more than 3 months ago. However, evidence from this same RCT suggests benefits do not extend to anxiety symptoms. Moreover, moderate quality evidence from this RCT (N=150) suggests a significantly higher rate of discontinuation associated with family therapy.
- Very low quality single-RCT (N=30-34) evidence suggests a delayed but large and statistically significant benefit of parent training (CBT with parent-only) relative to TAU on improving clinician-rated PTSD symptomatology and emotional and behavioural problems at 2-year follow-up (effects non-significant at endpoint and 3-, 6- and 12-month follow-up) for children and young people with PTSD who were exposed to trauma more than 3 months ago. Very low quality evidence from the same RCT (N=35) suggests large and statistically significant benefits of CBT with the parent-only on improving children's depression symptoms at endpoint and 2-year follow-up, although effects are non-significant at 3-, 6- and 12-month follow-up. No evidence for discontinuation is available.
- Very low quality single-RCT (N=41) evidence suggests delayed moderate and statistically significant benefits of parent training (CBT with parent-only) relative to trauma-focused CBT on improving PTSD symptomatology and depression symptoms at 2-year follow-up, for children and young people with PTSD who were exposed to trauma more than 3 months ago. Conversely evidence from the same study (N=38) suggests benefits in favour of trauma-focused CBT relative to parent training on emotional and behavioural problems at 6- and 12-month follow-up (non-significant at endpoint, 3-month or 2-year follow-up). No evidence for discontinuation is available.
- • Very low quality single-RCT (N=24) evidence suggests non-significant differences between combined trauma-focused CBT for the child and parent training relative to trauma-focused CBT (for the child)-only on clinician-rated PTSD symptomatology, anxiety or depression symptoms, emotional and behavioural problems or global functioning (at endpoint and 3-month follow-up) for children and young people with PTSD who were exposed to trauma more than 3 months ago. Low quality evidence from this same RCT also suggests non-significant effects on discontinuation.

## 49 Play therapy for delayed treatment (>3 months)

 Very low quality single-RCT (N=129) evidence suggests large and statistically significant benefits of play therapy relative to TAU on improving self-rated PTSD

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- symptomatology and anxiety and depression symptoms for children and young people with PTSD who were exposed to trauma more than 3 months ago.

  Evidence from this same RCT (N=131) also suggests a lower rate of
- discontinuation may be associated with play therapy relative to TAU, however absolute numbers are small and this effect is not statistically significant.
- Very low quality single-RCT (N=26) evidence suggests a non-significant difference between play therapy and trauma-focused CBT on self-rated PTSD symptomatology for children and young people with PTSD who were exposed to trauma more than 3 months ago. Low quality evidence from this same RCT (N=31) suggests a lower rate of discontinuation may be associated with play therapy relative to trauma-focused CBT, however this effect is not statistically significant.

## 13 Economic evidence statements

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- Evidence from 1 Australian model-based study suggests that trauma-focused CBT alone or combined with SSRIs is more cost-effective than counselling. The study is partially applicable to the NICE decision-making context and is characterised by potentially serious limitations.
- Evidence from 1 Australian model-based study suggests that trauma-focused CBT combined with SSRIs is more cost-effective than trauma-focused CBT alone. The study is partially applicable to the NICE decision-making context and is characterised by potentially serious limitations.
- Evidence from 1 Australian model-based study suggests that trauma-focused CBT is more cost-effective than treatment as usual. The study is partially applicable to the NICE decision-making context and is characterised by potentially serious limitations.
- Evidence from 1 UK cost-minimisation analysis conducted alongside a RCT (N=75) suggests that individual psychodynamic psychotherapy was significantly costlier than psychoeducational group therapy. The study is partially applicable to the NICE decision-making context and is characterised by potentially serious limitations.
- Evidence from 1 UK cost-utility study that extrapolated efficacy and cost data from a RCT (N=29) suggests that trauma-focused cognitive therapy is more costeffective than waitlist. The study is partially applicable to the NICE decisionmaking context and is characterised by potentially serious limitations.
- Evidence from the guideline economic analysis suggests that individual forms of TF-CBT and, to a lesser degree, play therapy are cost-effective in the treatment of children and young people with PTSD. Family therapy and supportive counselling do not appear to be cost-effective relative to other interventions and, under some scenarios, supportive counselling is less cost-effective than no treatment. Inbetween, there is another group of interventions (EMDR, group CBT and parent training) with modest relative cost effectiveness, which is affected by the alternative scenarios tested. The economic analysis is directly applicable to the NICE decision-making context and is characterised by minor limitations.

#### 44 Recommendations

 Consider individual trauma-focused CBT for children aged under 7 years with a diagnosis of PTSD or clinically important symptoms of PTSD more than 1 month after a traumatic event.

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- Consider individual trauma-focused CBT for children aged 7 to 17 years
   within 1-3 months of a traumatic event with a diagnosis of PTSD or
   clinically important symptoms of PTSD.
- 4 3. Offer individual trauma-focused CBT to children and young people aged 7 to 17 years with a diagnosis of PTSD or clinically important symptoms of PTSD more than 3 months after a traumatic event.

## 4. Trauma-focused CBT for children and young people should:

- be based on a validated manual
  - typically be provided over 6–12 sessions
- be adapted to the child or young person's age and development
- involve parents or carers as appropriate
  - include psychoeducation about reactions to trauma, strategies for managing arousal and safety planning
  - involve elaboration and processing of the trauma memories
  - involve restructuring trauma-related meanings for the individual
    - provide help to overcome avoidance
    - prepare them for the end of treatment
    - include planning booster sessions if needed, particularly in relation to significant dates (for example trauma anniversaries).
  - Consider eye movement desensitisation and reprocessing (EMDR) for children and young people aged 7 to 17 years with a diagnosis of PTSD or clinically important symptoms of PTSD more than 3 months after a traumatic event only if they do not respond to or engage with traumafocused CBT.

## 25 Rationale and impact

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## 26 Why the committee made the recommendations

- 27 The evidence showed that trauma-focused CBT is effective in improving PTSD
- 28 symptoms and other important outcomes, and that improvements last for at least a
- 29 year. Benefits were seen for different specific trauma-focused interventions and
- different types of trauma. Trauma-focused CBT is more effective, as well as more
- 31 cost effective when it is provided individually than in a group so the committee
- 32 agreed it should be delivered individually.
- The evidence suggested that trauma-focused CBT was effective for children both
- over and under 7 years. Most of the evidence came from older children, so the
- committee could not recommend it with the same certainty for under 7s but agreed it
- should be thought of as an option for them.
- 37 There was no evidence for early treatment (within 1-3 months of a traumatic event)
- 38 with trauma-focused CBT relative to a non-active control so the committee could not
- recommend it with the same certainty as for more than 3 months after trauma.
- 40 However the committee extrapolated from the broad evidence base for benefits more
- 41 than 3 months after trauma and their clinical experience and agreed that trauma-
- 42 focused CBT should be an option for treatment within this earlier time period.

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- 1 Although specific trauma-focused CBT interventions use the same broad approach
- the committee was concerned that psychological interventions are not always
- delivered in a consistent way, so they agreed to specify the structure and content.
- 4 There was limited evidence for eye movement desensitisation and reprocessing
- 5 (EMDR) suggesting possible benefits on PTSD symptoms in children aged over 7
- 6 years. Based on uncertainties in this evidence the committee decided it should be
- 7 considered only if children do not respond to or engage with trauma-focused CBT, an
- 8 intervention that is supported by better evidence.

## 9 Impact of the recommendations on practice

- 10 The recommendations for trauma-focused CBT more than 3 months after a traumatic
- event are in line with previously recommended practice and the committee were not
- aware of wide variation in practice. Recommending trauma-focused CBT within 1-3
- months after a traumatic event and also recommending EMDR as an option are both
- new, but any impact on practice should be limited by the fact that these are weaker
- 15 ('consider') recommendations and that EMDR should only be considered for children
- who do not respond to or engage with trauma-focused CBT.
- 17 Previous treatment recommendations were made for children with PTSD, whereas
- current recommendations are relevant to children and young people with a diagnosis
- of PTSD or with clinically important symptoms of PTSD. The committee noted that
- 20 the structure, content and time of the assessment, as well as the benefits from
- 21 treatment are broadly the same for both populations (i.e. those diagnosed with PTSD
- and those identified as having clinically important symptoms of PTSD) and expressed
- 23 the view that the population covered in the current treatment recommendations does
- 24 not represent a significant broadening of the population that was covered by the
- 25 previous guideline recommendations; indeed many children and young people with
- 26 clinically important symptoms of PTSD that are below the diagnostic threshold for
- 27 PTSD will eventually develop a diagnosis of PTSD.

## 28 The committee's discussion of the evidence

## 29 Interpreting the evidence

#### 30 Outcomes that matter the most

- 31 Critical outcomes were measures of PTSD symptom improvement on validated
- 32 scales, remission (as defined as a loss of diagnosis or scoring below threshold on a
- validated scale), and response (as measured by an agreed percentage improvement
- in symptoms and/or by a dichotomous rating of much or very much improved).
- 35 Attrition from treatment (for any reason) was also considered an important outcome,
- as a proxy for the acceptability and/or tolerability of treatment. The committee
- 37 considered dissociative symptoms, personal/social/educational functioning (including
- 38 global functioning/functional impairment, sleeping difficulties, and quality of life), and
- 39 symptoms of a coexisting condition (including anxiety, depression and emotional and
- 40 behavioural problems) as important but not critical outcomes. This distinction was
- 41 based on the primacy of targeting the core PTSD symptoms, whilst acknowledging
- 42 that broader symptom measures may be indicators of a general pattern of effect.
- 43 Generally change scores were favoured over final scores as although in theory
- randomisation should balance out any differences at baseline, this assumption can
- 45 be violated by small sample sizes. The committee also expressed a general
- 46 preference for self-rated PTSD symptomatology (over clinician-rated or parent-rated
- 47 measures), however, in considering psychological interventions (relative to

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- 1 pharmacological interventions) a greater emphasis was placed on triangulating
- 2 effects on self-rated PTSD symptomatology with clinician-rated outcome measures,
- 3 given that the latter but not the former could be blinded.

## 4 The quality of the evidence

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- With the exception of a few outcomes of moderate quality, all the evidence reviewed
- 6 was of low or very low quality, reflecting the high risk of bias associated with the
- 7 studies (including for instance, high risk of bias associated with randomisation
- 8 method as reflected by significant group differences at baseline, and lack of/unclear
- 9 blinding of outcome assessment), the small numbers in many trials and the
- imprecision of many of the results (in terms of both the width of the confidence
- intervals and the failure to meet the optimal information size). However, the
- 12 committee agreed to make a strong recommendation despite uncertainty in the
- evidence, as the breadth of outcomes considered allowed triangulation of effects,
- and greater confidence was conferred where long-term follow-up was available.

## Consideration of clinical benefits and harms

- 16 The committee discussed the strength and breadth of the evidence for trauma-
- 17 focused CBT, with benefits observed on both clinician-rated and self-rated measures
- of PTSD symptomatology, the rate of remission and response, and on other
- 19 outcomes including depression, anxiety, emotional and behavioural problems, quality
- of life and global functioning. Although there was evidence for some differential
- 21 relative efficacy on clinician-rated PTSD symptomatology clinically important and
- statistically significant benefits were observed across: a range of trauma types
- 23 (including motor vehicle collisions, witnessing war as a civilian, sexual abuse and
- 24 mixed trauma types); both single and multiple incident index traumas; both those with
- a diagnosis of PTSD and those with clinically important symptoms (who may not
- 26 necessarily have a diagnosis); across specific trauma-focused intervention types
- 27 (both those that place emphasis on exposure and those that place emphasis on
- cognitive techniques); across formats (individual, caregiver and child, and group);
- and for both studies where the age range includes children aged 7 years and under,
- and where the age range only includes over 7s. Taken together with evidence
- 31 suggesting that benefits are potentially long-lasting, the committee agreed that
- 32 trauma-focused CBT should be offered as a first-line treatment to children and young
- people with PTSD. The committee discussed the evidence that showed a trend for a
- 34 higher rate of discontinuation with trauma-focused CBT relative to waitlist, TAU or no
- 35 treatment, and agreed that, given that this effect was not statistically significant and
- the comparison against supportive counselling showed a trend in favour of trauma-
- 37 focused CBT for lower discontinuation, the benefits of trauma-focused CBT
- 38 outweighed any potential harm.
- 39 The committee noted that although interventions within the trauma-focused CBT
- 40 class are using the same broad approach and there is considerable overlap in the
- 41 techniques and proposed mechanisms of the various versions of trauma-focused
- 42 CBT, this class is a somewhat broad umbrella and it was therefore important to
- specify the content and structure of the recommended intervention. The committee
- 44 also expressed concern that psychological interventions are not always implemented
- 45 consistently. For example, audits have suggested less-than-recommended number of
- 46 sessions are used in practice. The recommended structure and content of trauma-
- focused CBT (number of sessions, manualised, involvement of parents or carers, included content) is informed by the interventions in the RCT's included in the
- 49 evidence review, and modified by the expert advice of the committee. This
- recommendation seeks to ensure clarity and consistency, and that use in routine

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practice reflects the interventions in the clinical trials on which efficacy estimates are based.

3 The committee also considered the limited evidence suggesting non-significant 4 differences on PTSD outcomes between EMDR and trauma-focused CBT up to 12-5 month follow-up, and reflected that although this evidence base is not sufficiently 6 powered to detect non-inferiority, the lack of a significant difference suggests EMDR 7 may also have benefits for improving PTSD symptomatology. Evidence also 8 suggested that EMDR may be superior to trauma-focused CBT for emotional and 9 behavioural problems. The committee also discussed the limited evidence comparing 10 EMDR against waitlist or TAU which again was potentially indicative of benefit on 11 self-rated PTSD symptomatology although this effect was not statistically significant 12 and effects on clinician-rated PTSD symptomatology were neither clinically important 13 nor statistically significant. As in the head-to-head comparison, EMDR was found to 14 have more convincing benefits on other important outcomes, in this case both 15 emotional and behavioural problems and quality of life. Based on the uncertainty in 16 the evidence the committee judged a weak recommendation to be appropriate, and 17 based on the superior clinical efficacy of trauma-focused CBT the committee agreed 18 that EMDR should only be considered if children who do not respond to or engage 19 with trauma-focused CBT.

Given the considerable evidence base for trauma-focused CBT, the committee considered it appropriate to set a relatively high bar for other interventions. For some interventions (such as emotional freedom technique [EFT], combined trauma-focused CBT and parent training, child-parent psychotherapy using play, parent training, and family therapy), there is limited evidence for efficacy but the evidence base was considered too small to be confident that the benefits observed are true effects and thus a recommendation could not be supported. For other interventions, such as supportive counselling, the suggestion of inferiority to trauma-focused CBT, together with the non-significant effects relative to no treatment or waitlist, were sufficient for the committee to decide that a recommendation was not appropriate. Play therapy looked potentially more promising and required greater scrutiny and deliberation. However, given that less is known about the breadth of effects (no evidence for clinician-rated PTSD symptomatology, remission, other important associated symptoms and no follow-up) and there was some difficulty in pinpointing the core active ingredient of a play therapy (given differences between the two interventions in this category in terms of the inclusion of cognitive elements), the committee came to the decision that the evidence was not sufficient to warrant a recommendation at this time.

## 38 Cost effectiveness and resource use

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- Existing evidence suggests that trauma-focused CBT, alone or combined with SSRIs is a cost-effective option for the treatment of PTSD in children and young people,
- 41 compared with counselling, TAU or no treatment. Individual psychodynamic
- 42 psychotherapy appears to be less cost-effective than psychoeducational group
- therapy. The committee took existing economic evidence into account but noted that
- some of this evidence is only partially applicable to the UK as the studies were
- 45 conducted in other countries, it assesses the relative cost effectiveness of a limited
- 46 number of interventions, and is characterised by potentially serious limitations.
- The committee considered the results of the base-case guideline economic analysis
- 48 when making recommendations, which was informed by a NMA of overall good
- 49 quality, although some of the secondary analyses were characterised by a limited
- 50 evidence base at endpoint. Results of the guideline economic analysis were directly

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- applicable to the NICE decision-making context and were thus given more weight
- than existing evidence. The guideline base-case economic analysis was overall
- 3 characterised by minor limitations, so the committee were confident to use its
- 4 findings to support recommendations.
- 5 Results suggested that individual forms of trauma-focused CBT and, to a lesser
- 6 degree, play therapy are cost-effective in the treatment of children and young people
- 7 with PTSD. Family therapy and supportive counselling do not appear to be cost-
- 8 effective relative to other interventions and, under some scenarios, supportive
- 9 counselling is less cost-effective than no treatment. In-between, there is another
- 10 group of interventions (EMDR, group CBT and parent training) with modest relative
- 11 cost effectiveness, which is somewhat affected by the alternative scenarios tested.
- 12 Results were robust to scenarios tested through deterministic sensitivity analysis.
- 13 The committee noted that all forms of individual trauma-focused CBT were cost-
- 14 effective, although their relative cost effectiveness was slightly affected by the
- different scenarios and assumptions tested. This evidence, combined with the fact
- that trauma-focused CBT has the largest empirical database among all interventions
- 17 tested on children and young people with PTSD led the committee to make a
- 18 recommendation for individual trauma-focused CBT.
- 19 Play therapy was also shown to be cost-effective in economic analysis, however
- 20 results were based on a limited evidence base (2 trials). As already reported, the
- 21 committee had some difficulty in pinpointing the core active ingredient of a play
- therapy (given differences between the two interventions in this category in terms of
- the inclusion of cognitive elements). Moreover, the committee noted that the resource
- use associated with the interventions differed considerably between the 2 trials,
- suggesting a less well-defined intervention, thus introducing uncertainty in the results
- of the economic analysis. For this reason and because of the limited evidence base
- the committee were reluctant to make a recommendation for play therapy.
- 28 Of the remaining interventions, EMDR, group CBT and parent training showed a
- 29 modest relative cost effectiveness. The committee considered the cost effectiveness
- of EMDR alongside the clinical evidence base and decided to make a 'consider'
- 31 recommendation for children and young people aged 7-18 years who do not respond
- 32 to or engage with trauma-focused CBT.
- The committee did not make any recommendation for group CBT, as it is less cost-
- 34 effective than individual trauma-focused CBT, individual trauma-focused CBT was
- already recommended as a first-line option, so no further benefits were expected to
- be gained by a potential recommendation on group CBT.
- 37 The committee did not make a recommendation on parent training, despite its
- 38 modest relative cost effectiveness, because this result was based on a very limited
- 39 evidence base (N=49).
- The committee anticipated that the recommendations will result in a moderate
- 41 change in practice. The only strong ('offer') recommendation for trauma-focused CBT
- more than 3 months after a traumatic event was also a strong recommendation in the
- previous guideline and the committee were not aware of wide variation in practice.
- The recommendations for trauma-focused CBT within 1-3 months after a traumatic
- 45 event and the recommendation for EMDR are new. However, changes in practice will
- be limited by the fact that these are weaker recommendations and, in the case of
- 47 EMDR, the recommendation should be considered if children do not respond to or
- 48 engage with trauma-focused CBT. Moreover, it is anticipated that children with a
- 49 diagnosis of PTSD or clinically important symptoms of PTSD within 1-3 months after

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- 1 a traumatic event, as well as children who do not respond to or engage with trauma-
- 2 focused CBT, may be currently accessing services and receiving alternative
- 3 interventions of lesser known effectiveness, which are anticipated to be replaced, at
- 4 least partly, by the recommended interventions, and therefore implementation costs
- of newly recommended interventions are expected to be offset, to some extent, by
- 6 cost-savings of interventions forgone.
- When assessing the impact of treatment recommendations on available resources,
- 8 the committee was aware that previous recommendations were made for children
- 9 with PTSD, whereas current recommendations are relevant to children with a
- diagnosis of PTSD or with clinically important symptoms of PTSD. The latter are
- identified when people score above a pre-determined threshold on a validated PTSD
- 12 symptom scale, which is indicative but not confirmatory of a diagnosis of PTSD. The
- 13 committee noted that the assessment of a person with suspected PTSD includes a
- general assessment of mental state, specific questions about the traumatic event(s),
- 15 enquiries into specific traumatic hyper vigilance and intrusive thoughts and
- assessment of the impact of the symptoms on personal and social functioning. In
- 17 current practice, the structure, content and time of the assessment is the same for
- 18 people for whom a diagnosis of PTSD has been made and for people who have been
- 19 identified to experience clinically important symptoms of PTSD. The committee noted
- 20 that the decision to start treatment in both populations is influenced by the severity of
- 21 symptoms, the trajectory of symptoms, any co-morbid mental disorders and the
- 22 individual's preference for treatment. The committee expressed the opinion that the
- 23 impact of experiencing clinically important PTSD symptoms on the person's social
- and personal functioning may be broadly similar to the impact of a formal diagnosis
- of PTSD, depending on the presence and/or intensity of the factors described above
- 26 (i.e. severity and trajectory of symptoms and any co-morbid mental disorders) and
- decided that treatment recommendations should focus on both populations. The
- committee expressed the view that the population of children and young people
- 29 covered in the current treatment recommendations does not represent a significant
- 30 broadening of the population that was covered by the previous guideline
- 31 recommendations, and indeed many children and young people with clinically
- 32 important symptoms of PTSD that are below the diagnostic threshold for PTSD will
- eventually develop a diagnosis of PTSD.

## 34 Other factors the committee took into account

- 35 The committee discussed the inclusion of family members in the treatment of children
- and young people and concluded that the carers of the child or young person should
- 37 be included and involved in the treatment as and when appropriate and deemed to
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## 10 Combined somatic and cognitive therapies

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## 17 **Deblinger 1996/1999**

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# 1 Psychosocial interventions for the treatment of PTSD in

# 2 children and young people

## 3 Introduction

- 4 Psychosocial interventions will be considered as classes of intervention (art therapy;
- 5 music therapy; meditation; psychoeducational interventions; peer support) and form
- 6 the subsections below.
- 7 Evidence for interventions in the following classes was also searched for but none
- 8 was found: mindfulness-based stress reduction (MBSR); nature-assisted therapies;
- 9 supported employment; practical support.

## 10 Art therapy: clinical evidence

#### 11 Included studies

- 12 Four studies of art therapy for the treatment of PTSD in children and young people
- were identified for full-text review. Of these 4 studies, 1 RCT (N=77) was included in
- 14 a single comparison.
- 15 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- symptoms, there were no included studies.
- 17 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 18 PTSD symptoms, 1 RCT (N=77) compared art therapy (in addition to TAU) with
- 19 attention-placebo (in addition to TAU) (Lyshak-Stelzer 2007).

## 20 Excluded studies

- 21 Three studies were reviewed at full text and excluded from this review due to non-
- 22 randomised group assignment, small sample size (N<10 per arm), or because
- 23 efficacy or safety data could not be extracted.
- 24 Studies not included in this review with reasons for their exclusions are provided in
- 25 Appendix K.

33

34

## 26 Summary of clinical studies included in the evidence review

- 27 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 28 study evidence tables in Appendix D.
- 29 Table 31 provides a brief summary of the included study and evidence from this
- 30 study is summarised in the clinical GRADE evidence profile below (Table 32).
- 31 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 32 study evidence tables in Appendix D.

# Table 31: Summary of included studies: Art therapy for delayed treatment (>3 months)

| Comparison                          | Art therapy (+ TAU) versus attention-placebo (+ TAU) |
|-------------------------------------|--|
| Total no. of studies (N randomised) | 1 (77)   |
| Study ID                            | Lyshak-Stelzer 2007                                  |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Comparison                               | Art therapy (+ TAU) versus attention-placebo (+ TAU)  |
|--|---|
| Country                                  | US  |
| Diagnostic status                        | Clinically important PTSD symptoms (scoring above a threshold on validated scale)   |
| Mean months since onset of PTSD          | NR  |
| Mean age (range)                         | 15.1 (13-17)  |
| Sex (% female)                           | 45  |
| Ethnicity (% BME)                        | 82  |
| Coexisting conditions                    | NR  |
| Mean months since traumatic event        | NR  |
| Type of traumatic event                  | Mixed: Frequently reported trauma types included: Physically abused or threatened with physical abuse at home (62%); Witnessing physical abuse at home (50%); Being in a bad accident (50%); Witnessing shooting, beating, or threats in neighbourhood (47%); Sexual abuse (46%); Beaten, shot at, or threatened in neighbourhood (45%); Serious medical problem (40%); Being in a disaster (weather, fire, etc.) (19%) |
| Single or multiple incident index trauma | Multiple  |
| Lifetime experience of trauma            | NR  |
| Intervention details                     | Trauma-focused expressive art therapy (+ TAU). Each participant completed collages or drawings to express a narrative of his/her "life story"   |
| Intervention format                      | Group   |
| Intervention intensity                   | 16x weekly sessions (length of session NR)  |
| Comparator                               | Attention-placebo (+ TAU): standard arts and craft activity group already in use at the two facilities  |
| Intervention length (weeks)              | 16  |

- 1 BME –Black and minority ethnic; NR-Not reported; PTSD-Post-traumatic stress disorder; TAU-
- 1 BME –Black and mir 2 Treatment as usual.
- 3 See appendix D for full evidence tables.

## 4 Quality assessment of clinical studies included in the evidence review

- 5 The clinical evidence profile for this review (art therapy for the treatment of PTSD in
- 6 children and young people) are presented in Table 32.

2

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## Table 32: Summary clinical evidence profile: Art therapy (+ TAU) versus attention-placebo (+ TAU) for delayed treatment (>3 months)

|  |  | ( 1710) for dolay   |                                |  |  |
|--|--|---|--------------------------------|--|--|
|  | Illustrative comparative risks* (95% CI)                 |   |                                |  |  |
| Outcomes   | Assum<br>ed risk<br>Attentio<br>n-<br>placebo<br>(+ TAU) | Corresponding risk Art therapy (+ TAU)  | Relative<br>effect<br>(95% CI) | No of<br>Participant<br>s<br>(studies) | Quality<br>of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatology<br>clinician-rated<br>UCLA PTSD-RI<br>administered via<br>structured<br>interview<br>format; change<br>score<br>Follow-up:<br>mean 16 weeks |  | The mean PTSD symptomatology clinician-rated in the intervention groups was 1.79 standard deviations lower (2.67 to 0.91 lower) |                                | 29<br>(1 study)                        | low <sup>1,2</sup>                       |

- CI=confidence interval; PTSD=post-traumatic stress disorder; SMD=standardised mean difference;
- TAU=treatment as usual; UCLA PTSD-RI= UCLA PTSD-Reaction Index
  - <sup>1</sup> OIS not met (N<400)
- 3 4 5 6 7 <sup>2</sup> Data is not reported/cannot be extracted for all outcomes and this is interim report but unable to locate full report
- 8 See appendix F for full GRADE tables.

## 9 Music therapy: clinical evidence

#### 10 Included studies

- 11 One study of music therapy for the treatment of PTSD in children and young people
- was identified for full-text review. This study could not be included. 12

## 13 Excluded studies

- 14 One study was reviewed at full text and excluded from this review due to non-
- 15 randomised group assignment.
- 16 Studies not included in this review with reasons for their exclusions are provided in
- 17 Appendix K.

## 18 Meditation: clinical evidence

#### 19 Included studies

- 20 Five studies of meditation for the treatment of PTSD in children and young people
- were identified for full-text review. Of these 5 studies, 1 RCT (N=82) was included in 21
- 22 a single comparison for meditation.
- 23 For early treatment (intervention initiated 1-3 months post-trauma) of PTSD
- 24 symptoms, there were no included studies.

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

- 1 For delayed treatment (intervention initiated more than 3 months post-trauma) of
- 2 PTSD symptoms, 1 RCT (N=82) compared meditation with waitlist (Gordon
- 3 2006/Gordon
- 4 2008 [protocol and published paper]).

## 5 Excluded studies

- 6 Four studies were reviewed at full text and excluded from this review because the
- 7 population was outside scope (<80% of participants are eligible for the review and
- 8 disaggregated data cannot be obtained), the study was unpublished (registered on
- 9 clinical trials.gov and author contacted for full trial report but not provided), or the
- paper was a systematic review with no new useable data and any meta-analysis
- 11 results not appropriate to extract.
- 12 Studies not included in this review with reasons for their exclusions are provided in
- 13 Appendix K.

21

22

## 14 Summary of clinical studies included in the evidence review

- 15 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 16 study evidence tables in Appendix D.
- 17 Table 33 provides a brief summary of the included study and evidence from this
- study is summarised in the clinical GRADE evidence profile below (Table 34).
- 19 See also the study selection flow chart in Appendix C, forest plots in Appendix E and
- 20 study evidence tables in Appendix D.

# Table 33: Summary of included studies: Meditation for delayed treatment (>3 months)

| 1110111115)                              |  |
|--|--|
| Comparison                               | Meditation versus waitlist   |
| Total no. of studies (N randomised)      | 1 (82)   |
| Study ID                                 | Gordon 2006/2008   |
| Country                                  | Kosovo   |
| Diagnostic status                        | PTSD diagnosis according to ICD/DSM criteria   |
| Mean months since onset of PTSD          | NR   |
| Mean age (range)                         | 16.3 (14-18)   |
| Sex (% female)                           | 76   |
| Ethnicity (% BME)                        | NR   |
| Coexisting conditions                    | NR   |
| Mean months since traumatic event        | NR   |
| Type of traumatic event                  | Witnessing war as a civilian: Kosovar adolescents  |
| Single or multiple incident index trauma | Multiple   |
| Lifetime experience of trauma            | NR   |
| Intervention details                     | Mind-body skills group, combines meditation with spoken and written word exercises, drawing and movement in a small-group school setting |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Comparison                  | Meditation versus waitlist                  |
|-----------------------------|---|
| Intervention format         | Group                                       |
| Intervention intensity      | 12x twice-weekly 2-hour sessions (24 hours) |
| Comparator                  | Waitlist                                    |
| Intervention length (weeks) | 6   |

- 1 BME -Black and minority ethnic; NR-Not reported; PTSD-Post-traumatic stress disorder.
- 2 See appendix D for full evidence tables.

## 3 Quality assessment of clinical studies included in the evidence review

- 4 The clinical evidence profile for this review (meditation for the treatment of PTSD in
- children and young people) is presented in Table 34. 5

## Table 34: Summary clinical evidence profile: Meditation versus waitlist for delayed treatment (>3 months)

|   |                             | •  |                             |                                    |                               |
|---|-----------------------------|--|-----------------------------|------------------------------------|-------------------------------|
|   | Illustrative co<br>(95% CI) | omparative risks*  | Relativ                     |                                    | Quality                       |
| Outcomes  | Assumed risk Waitlist       | Corresponding risk Meditation  | e effect<br>(95%<br>CI)     | No of<br>Participants<br>(studies) | of the<br>evidence<br>(GRADE) |
| PTSD<br>symptomatology<br>self-rated<br>HTQ change<br>score<br>Follow-up:<br>mean 6 weeks                         | Wallist                     | The mean PTSD symptomatology self-rated in the intervention groups was 1.65 standard deviations lower (2.17 to 1.13 lower) | Gij                         | 77<br>(1 study)                    | low <sup>1,2</sup>            |
| Discontinuation<br>Number of<br>participants lost<br>to follow-up for<br>any reason<br>Follow-up:<br>mean 6 weeks | 49 per 1000                 | 73 per 1000<br>(13 to 415)   | RR 1.5<br>(0.26 to<br>8.51) | 82<br>(1 study)                    | low <sup>3</sup>              |

- CI=confidence interval; HTQ= Harvard Trauma Questionnaire; PTSD=post-traumatic stress disorder;
- 8 9 10 RR=risk ratio; SMD=standardised mean difference
- <sup>1</sup> Risk of bias is high or unclear across multiple domains
- 11 <sup>2</sup> OIS not met (N<400)

6

- 12 <sup>3</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm
- 13 See appendix F for full GRADE tables.

## 14 Psychoeducational interventions: clinical evidence

#### 15 Included studies

- 16 Seven studies of psychoeducational interventions for the treatment of PTSD in
- 17 children and young people were identified for full-text review. None of these studies
- could be included 18

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

#### 1 Excluded studies

- 2 Seven studies were reviewed at full text and excluded from this review. Reasons for
- 3 exclusion included non-randomised group assignment, non-validated outcome
- 4 measures and small sample size (N<10 per arm).
- 5 Studies not included in this review with reasons for their exclusions are provided in
- 6 Appendix K.

## 7 Peer support: clinical evidence

#### 8 Included studies

- 9 Four studies of peer support for the treatment of PTSD in children and young people
- were identified for full-text review. None of these studies could be included.

#### 11 Excluded studies

- 12 Four studies were reviewed at full text and excluded from this review because the
- intervention was not targeted at PTSD symptoms, or efficacy or safety data could not
- 14 be extracted.
- 15 Studies not included in this review with reasons for their exclusions are provided in
- 16 Appendix K.

#### 17 Economic evidence

#### 18 Included studies

- 19 The systematic search of the literature did not identify any economic studies
- 20 assessing the cost effectiveness of psychosocial interventions for the treatment of
- 21 PTSD in children and young people. The search strategy for economic studies is
- 22 provided in Appendix B.

## 23 Excluded studies

- 24 No economic studies of psychosocial interventions for the treatment of PTSD in
- 25 children and young people were reviewed at full text and excluded.

## 26 Economic model

- 27 No economic modelling was undertaken in this area because other topics were
- agreed as higher priorities for economic evaluation.

## 29 Resource impact

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- 30 As no recommendations were made in this area and psychosocial interventions for
- 31 the treatment of PTSD in children and young people are not in widespread use in
- 32 routine clinical practice, there is no impact on resources.

## 33 Clinical evidence statements

## 34 Art therapy for delayed treatment (>3 months)

 Low quality single-RCT (N=29) evidence suggests a large and statistically significant benefit of art therapy (in addition to TAU) relative to attention-placebo

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

1 (in addition to TAU) on improving clinician-rated PTSD symptomatology for children with PTSD who were exposed to trauma more than 3 months ago. No

3 other outcomes are available for art therapy.

## 4 Meditation for delayed treatment (>3 months)

 Low quality single-RCT (N=77) evidence suggests a large and statistically significant benefit of meditation relative to waitlist on improving self-rated PTSD symptomatology for children with PTSD who were exposed to trauma more than 3 months ago. Evidence from this same RCT (N=82) suggests a trend for a higher rate of discontinuation associated with meditation relative to waitlist, however absolute differences are small and this effect is not statistically significant.

## 11 Economic evidence statements

- 12 No economic evidence on psychosocial interventions for the treatment of PTSD in
- 13 children and young people was identified and no primary economic modelling was
- 14 undertaken.

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#### 15 Recommendations

- 16 No recommendations were made for psychosocial interventions for the treatment of
- 17 PTSD in children and young people.

## 18 Rationale and impact

## 19 Why the committee didn't make any recommendations

- 20 Limited evidence showed that meditation and art therapy might have some benefit on
- 21 PTSD symptoms in children and young people. However, because there were too
- 22 many uncertainties in the evidence and there was much better evidence supporting
- trauma-focused CBT, the committee did not make any recommendations.

#### 24 The committee's discussion of the evidence

## 25 Interpreting the evidence

## 26 Outcomes that matter the most

- 27 Critical outcomes were measures of PTSD symptom improvement on validated
- scales, remission (as defined as a loss of diagnosis or scoring below threshold on a
- validated scale), and response (as measured by an agreed percentage improvement
- in symptoms and/or by a dichotomous rating of much or very much improved).
- 31 Attrition from treatment (for any reason) was also considered an important outcome,
- 32 as a proxy for the acceptability and/or tolerability of treatment. The committee
- 33 considered dissociative symptoms, personal/social/educational functioning (including
- global functioning/functional impairment, sleeping difficulties, and quality of life), and
- 35 symptoms of a coexisting condition (including anxiety, depression and emotional and
- 36 behavioural problems) as important but not critical outcomes. This distinction was
- based on the primacy of targeting the core PTSD symptoms, whilst acknowledging that broader symptom measures may be indicators of a general pattern of effect.
- 39 Generally change scores were favoured over final scores as although in theory
- 40 randomisation should balance out any differences at baseline, this assumption can
- 41 be violated by small sample sizes. The committee also expressed a general
- 42 preference for self-rated PTSD symptomatology, however, in considering

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

- 1 psychosocial interventions (relative to pharmacological interventions) a greater
- 2 emphasis was placed on triangulating effects on self-rated PTSD symptomatology
- 3 with clinician-rated outcome measures, given that the latter but not the former could
- 4 be blinded.

## The quality of the evidence

- 6 All the evidence reviewed was of low quality, reflecting the high risk of bias
- 7 associated with the studies (including for instance, non-blind outcome assessment),
- 8 the small numbers in trials and the imprecision of many of the results (in terms of the
- 9 failure to meet the optimal information size). This uncertainty of the evidence is
- 10 reflected in the Committee decision to not make any recommendations for
- psychosocial interventions for the treatment of PTSD in children and young people.

## 12 Consideration of clinical benefits and harms

- 13 The Committee considered the evidence suggesting potential benefits of meditation
- or art therapy on improving self-rated or clinician-rated PTSD symptomatology
- 15 respectively. However, evidence for both interventions was limited to small single
- 16 studies, there was no evidence for effects on important associated symptoms or a
- 17 triangulation of effects on other PTSD outcomes, there was no evidence for
- discontinuation for art therapy, and a non-statistically significant trend for a higher
- 19 rate of discontinuation associated with meditation. Based on this uncertainty in the
- 20 evidence, the Committee were not confident in the robustness of the benefits
- 21 themselves or whether any benefits would outweigh any potential harms, and thus a
- 22 recommendation was not warranted.

#### 23 Cost effectiveness and resource use

- No evidence on the cost effectiveness of psychosocial interventions for the treatment
- of PTSD in children and young people was identified and no economic modelling was
- undertaken in this area. As there was very limited evidence of clinical benefit
- associated with psychosocial interventions, no recommendation was made. None of
- these interventions are in widespread use in routine clinical practice, therefore no
- 29 impact on resources is expected.

## 30 References for included studies

## 31 Art therapy

## 32 Lyshak-Stelzer 2007

- 33 Lyshak-Stelzer F, Singer P, Patricia SJ and Chemtob CM (2007) Art therapy for
- 34 adolescents with posttraumatic stress disorder symptoms: A pilot study. Art Therapy
- 35 24(4), 163-9

#### 36 Meditation

## 37 Gordon 2006/2008

- 38 Gordon JS (2006) Treatment of Posttraumatic Stress Disorder in Kosovar High
- 39 School Students Using Mind-Body Skills Groups: A Randomized Controlled Trial
- 40 [NCT00136357]. Available from:
- 41 https://clinicaltrials.gov/ct2/show/study/NCT00136357 [accessed 29.04.17]

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

- 1 Gordon JS, Staples JK, Blyta A, et al. (2008) Treatment of posttraumatic stress
- 2 disorder in postwar Kosovar adolescents using mind-body skills groups: a
- 3 randomized controlled trial. The Journal of clinical psychiatry 69(9), 1469-76

## 4 Other non-pharmacological interventions for the

## 5 treatment of PTSD in children and young people

## 6 Introduction

- 7 Potentially relevant evidence was only identified for one class of non-
- 8 pharmacological intervention, massage (see subsection below).
- 9 Evidence for interventions in the following classes was also searched for but none
- 10 was found: acupuncture; exercise; repetitive transcranial magnetic stimulation
- 11 [rTMS]; yoga).

## 12 Massage: clinical evidence

#### 13 Included studies

- One study of massage for the treatment of PTSD in children and young people was
- identified for full-text review. This study could not be included.

## 16 Excluded studies

- 17 One study was reviewed at full text and excluded from this review because efficacy
- or safety data could not be extracted.
- 19 Studies not included in this review with reasons for their exclusions are provided in
- 20 Appendix K.

## 21 Economic evidence

## 22 Included studies

- 23 The systematic search of the literature did not identify any economic studies
- 24 assessing the cost effectiveness of other non-pharmacological interventions for the
- 25 treatment of PTSD in children and young people. The search strategy for economic
- 26 studies is provided in Appendix B.

#### 27 Excluded studies

- 28 No economic studies of other non-pharmacological interventions for the treatment of
- 29 PTSD in children and young people were reviewed at full text and excluded.

## 30 Economic model

- 31 No economic modelling was undertaken in this area because other topics were
- 32 agreed as higher priorities for economic evaluation.

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## 1 Resource impact

- 2 As no recommendations were made in this area and other non-pharmacological
- 3 interventions for the treatment of PTSD in children and young people are not in
- 4 widespread use in routine clinical practice, there is no impact on resources.

## 5 Clinical evidence statements

- 6 No clinical evidence on other non-pharmacological interventions for the treatment of
- 7 PTSD in children and young people met inclusion criteria for this review.

## 8 Economic evidence statements

- 9 No economic evidence on other non-pharmacological interventions for the treatment
- of PTSD in children and young people was identified and no primary economic
- 11 modelling was undertaken.

## 12 Recommendations

- 13 No recommendations were made for other non-pharmacological interventions for the
- treatment of PTSD in children and young people.

## 15 Rationale and impact

## 16 Why the committee didn't make any recommendations

- 17 No evidence was found on other non-pharmacological interventions so the committee
- 18 did not make any recommendations.

## 19 The committee's discussion of the evidence

- 20 Interpreting the evidence
- 21 Outcomes that matter the most
- 22 No evidence was included in this review.
- 23 The quality of the evidence
- 24 No evidence was included in this review.
- 25 Consideration of clinical benefits and harms
- 26 No evidence was included in this review.

#### 27 Cost effectiveness and resource use

- 28 No evidence on the cost effectiveness of other non-pharmacological
- 29 interventions for the treatment of PTSD in children and young people
- was identified and no economic modelling was undertaken in this area.
- 31 As there was no clinical evidence available, no recommendation was
- 32 made. None of these interventions are in widespread use in routine

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

## 1 clinical practice, therefore no impact on resources is expected. Other

## 2 references

- 3 Alisic 2014
- 4 Alisic E, Zalta AK, Van Wesel F, et al. (2014) Rates of post-traumatic stress disorder
- 5 in trauma-exposed children and adolescents: meta-analysis. The British Journal of
- 6 Psychiatry 204(5), 335-340
- 7 Hiller **2016**
- 8 Hiller RM, MeiserStedman R, Fearon P, et al. (2016) Research Review: Changes in
- 9 the prevalence and symptom severity of child post- traumatic stress disorder in the
- 10 year following trauma-a meta- analytic study. Journal of Child Psychology and
- 11 Psychiatry 57(8), 884-898
- 12

# **Appendices**

# **Appendix A – Review protocols**

Review protocol for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

| Topic              | Pharmacological interventions for the treatment of PTSD in adults   |
|--------------------|---|
| Review question(s) | RQ. 1.2 For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?   |
| Sub-question(s)    | Where evidence exists, consideration will be given to the specific needs of: women who have been exposed to sexual abuse or assault, or domestic violence lesbian, gay, bisexual, transsexual or transgender people people from black and minority ethnic groups people who are homeless or in insecure accommodation asylum seekers or refugees or other immigrants who are entitled to NHS treatment people who have been trafficked people who are socially isolated (and who are not captured by any other subgroup listed) people with complex PTSD people with neurodevelopmental disorders (including autism) people with coexisting conditions (drug and alcohol misuse, common mental health disorders, eating disorders, personality disorders, acquired brain injury, physical disabilities and sensory impairments) people who are critically ill or injured (for instance after a vehicle crash) |
| Objectives         | To identify the most effective psychological, psychosocial or other non-pharmacological interventions for the treatment of PTSD in children and young people  |

| Topic        |  |
|--------------|--|
|              | Pharmacological interventions for the treatment of PTSD in adults  |
| Population   | Children and young people (aged under 18 years) with clinically important post-traumatic stress symptoms (more than one month after a traumatic event), defined by a diagnosis of PTSD according to DSM, ICD or similar criteria (including PTSD for children 6 years and younger) or clinically-significant PTSD symptoms as indicated by baseline scores above threshold on a validated scale (see PTSD scales listed under outcomes). |
|              | For mixed adult and children populations, where possible disaggregated data will be obtained. If this is not possible then the study will be categorised according to the mean age of the population (<18 years as children and young people and ≥18 years as adult).  |
|              | If some, but not all, of a study's participants are eligible for the review, where possible disaggregated data will be obtained. If this is not possible then the study will be included if at least 80% of its participants are eligible for this review.   |
| Exclude      | Trials of people with adjustment disorders   |
|              | Trials of people with traumatic grief  |
|              | Trials of people with psychosis as a coexisting condition  |
|              | Trials of people with learning disabilities  |
|              | Trials of women with PTSD during pregnancy or in the first year following childbirth   |
|              | Trials of adults in contact with the criminal justice system (not solely as a result of being a witness or victim)   |
| Intervention | Psychological interventions (psychological interventions listed below are examples of interventions which may be included either alone or in combination and delivered to the child or young person and/or a parent or carer in an individual or group format):  |
|              | Trauma-focused cognitive behavioural therapies (CBT), including cognitive therapy, cognitive processing therapy, compassion focused therapy, exposure therapy/prolonged exposure (PE), virtual reality exposure therapy (VRET), imagery rehearsal therapy, mindfulness-based cognitive therapy (MBCT) and narrative exposure therapy for traumatized children and adolescents (KidNET)   |
|              | Non-trauma-focused CBT, including stress inoculation training (SIT)  |
|              | Psychologically-focused debriefing (including single session debriefing)   |
|              | Eye movement desensitisation and reprocessing (EMDR)   |

| Topic |   |
|-------|---|
|       | Pharmacological interventions for the treatment of PTSD in adults   |
|       | Hypnotherapy  |
|       | Psychodynamic therapies, including traumatic incident reduction (TIR)   |
|       | Counselling, including non-directive/supportive/person-centred counselling  |
|       | Human givens therapy  |
|       | Combined somatic and cognitive therapies, including thought field therapy (TFT) and emotional freedom technique (EFT)   |
|       | Parent training/family interventions, including behavioural family therapy (such as Child and Family Traumatic Stress Intervention [CFTSI])   |
|       | Play therapy  |
|       | Psychosocial interventions (psychosocial interventions listed below are examples of interventions which may be included either alone or in combination):  Meditation                |
|       | Mindfulness-based stress reduction (MBSR)   |
|       | Nature-assisted therapies (including ecotherapy, horticultural therapy, therapeutic horticulture and nature-based therapy)  |
|       | Supported employment (including individual placement and support [IPS] supported employment and Veterans Health Administration Vocational Rehabilitation Programme [VRP])           |
|       | Practical support (including financial and housing)   |
|       | Psychoeducational interventions   |
|       | Peer support (including self-help groups and support groups)  |
|       | Other non-pharmacological interventions (other non-pharmacological interventions listed below are examples of   |
|       | interventions which may be included either alone or in combination):  |
|       | Acupuncture (including classical acupuncture, electroacupuncture, auricular acupuncture, laser acupuncture and acupoint stimulation [such as acupressure, moxibustion and tapping]) |

| Pharmacological interventions for the treatment of PTSD in adults   |
|---|
| Exercise (including anaerobic [such as heavy weight training, sprinting, high-intensity interval training] and aerobic [such as running/jogging, swimming, cycling and walking] exercise, both supervised and unsupervised) Repetitive transcranial magnetic stimulation (rTMS) Yoga (including all types of yoga)  |
| Combination interventions, such as combined psychological plus pharmacological versus pharmacological alone, will also be considered here.  |
| A distinction will be made between early interventions (delivered within 3 months of the traumatic event) and delayed interventions (delivered more than 3 months after the traumatic event)  |
| Exclude: Inoculation interventions for people who may be at risk of experiencing but have not experienced, a traumatic event Interventions that are not targeted at PTSD symptoms   |
| Any other intervention Treatment as usual Waitlist Placebo  |
| Efficacy PTSD symptomology (mean endpoint score or change in PTSD score from baseline) Diagnosis of PTSD (number of people meeting diagnostic criteria for PTSD according to DSM, ICD or similar criteria) Recovery from PTSD/Remission (number of people no longer meeting diagnostic criteria for PTSD according to DSM, ICD or similar criteria at endpoint, or endpoint scores below threshold on a validated scale) Response (as measured by an agreed percentage improvement in symptoms and/or by a dichotomous rating of much or very much improved on Clinical Global Impressions [CGI] scale)Relapse (number of people who remitted |
|   |

| Topic                                |  |
|--------------------------------------|--|
|                                      | Pharmacological interventions for the treatment of PTSD in adults  |
|                                      | at endpoint but at follow-up either met diagnostic criteria for PTSD according to DSM, ICD or similar criteria, or whose follow-up scores were above threshold on a validated scale) |
|                                      | The following PTSD scales will be included:  |
|                                      | Assessor-rated PTSD symptom scales   |
|                                      | Clinician-Administered PTSD Scale for Children and Adolescents for DSM–IV (CAPS–CA) or DSM-V (CAPS-CA-5)   |
|                                      | Anxiety Disorders Interview Schedule for Children for DSM-IV (ADIS-C)  |
|                                      | Schedule for Affective Disorders and Schizophrenia for School Age Children (K–SADS)  |
|                                      | Children's PTSD Inventory (CPTSDI)   |
|                                      | Self-report (parent-report) instruments of PTSD symptoms:  |
|                                      | Children's Impact of Event Scale/Children's Revised Impact of Event Scale (CRIES)  |
|                                      | Child Post Traumatic Stress Reaction Index (CPTS–RI)/UCLA PTSD Index for DSM-IV (UPID)/ CPTS-RI Revision 2 (also referred to as the PTSD Index for DSM-IV)                           |
|                                      | Child PTSD Symptom Scale (CPSS)  |
|                                      | Trauma Screening Checklist for Children (TSCC)   |
|                                      | Children's Reaction to Traumatic Events Scale (CRTES)  |
|                                      | Angie/ Andy Cartoon Trauma Scales (ACTS)/ Angie/Andy Parent Rating Scales  |
|                                      | Pediatric Emotional Distress Scale (PEDS)  |
|                                      | Acceptability/tolerability   |
|                                      | Acceptability of the intervention  |
|                                      | Discontinuation due to adverse effects   |
|                                      | Discontinuation due to any reason (including adverse effects)  |
| Important, but not critical outcomes | Dissociative symptoms as assessed with a validated scale including:  |
|                                      | Assessor-rated scales:   |
|                                      | Dissociation symptom cluster score on CAPS-CA  |

| Topic                     |  |
|---------------------------|--|
|                           | Pharmacological interventions for the treatment of PTSD in adults  |
|                           | Self-report (parent-report) scales:  |
|                           | Adolescent Dissociative Experiences Scale (A-DES)  |
|                           | Child Dissociative Checklist (CDC)   |
|                           | Personal, social, educational and occupational functioning   |
|                           | Emotional and behavioural problems (as assessed with a validated scale including Strengths and Difficulties Questionnaire [SDQ])   |
|                           | Sleeping difficulties (as assessed with a validated scale including Children's Sleep Habits Questionnaire [CSHQ], Sleep Disturbance Scale for Children [SDSC])   |
|                           | School attendance  |
|                           | Employment (for instance, number in paid employment)   |
|                           | Housing (for instance, number homeless or in insecure accommodation)   |
|                           | Quality of life (as assessed with a validated scale including Pediatric Quality of Life Inventory [PedsQL] and Warwick-Edinburgh Mental Well-being Scale [WEMWBS])   |
|                           | Coexisting conditions (note that target of intervention should be PTSD symptoms):  |
|                           | Symptoms of and recovery from a coexisting condition   |
|                           | Self-harm  |
|                           | Suicide  |
| Study design              | Systematic reviews of RCTs RCTs  |
| Include unpublished data? | Clinical trial registries (ISRCTN and ClinicalTrials.gov) will be searched to identify any relevant unpublished trials and authors will be contacted to request study reports (where these are not available online). Unpublished data will only be included where a full study report is available with sufficient detail to properly assess the risk of bias. Authors of unpublished evidence will be asked for permission to use such data, and will be informed that summary data from the study and the study's characteristics will be published in the full guideline |

| Topic                |   |
|----------------------|---|
|                      | Pharmacological interventions for the treatment of PTSD in adults   |
|                      | Conference abstracts and dissertations will not be included.  |
| Restriction by date? | All relevant studies from existing reviews from the 2005 guideline will be carried forward. No restriction on date for the updated search.  |
| Minimum sample size  | N = 10 in each arm  |
| Study setting        | Primary, secondary, tertiary, social care and community settings.   |
|                      | Treatment provided to troops on operational deployment or exercise will not be covered.   |
| The review strategy  | Reviews   |
|                      | If existing systematic reviews are found, the committee will assess their quality, completeness, and applicability to the NHS and to the scope of the guideline. If the committee agrees that a systematic review appropriately addresses a review question, a search for studies published since the review will be conducted.   |
|                      | Data Extraction (selection and coding)  |
|                      | Citations from each search will be downloaded into EndNote and duplicates removed. Titles and abstracts of identified studies will be screened by two reviewers for inclusion against criteria, until a good inter-rater reliability has been observed (percentage agreement =>90% or Kappa statistics, K>0.60). Initially 10% of references will be double-screened. If inter-rater agreement is good then the remaining references will be screened by one reviewer. All primary-level studies included after the first scan of citations will be acquired in full and re-evaluated for eligibility at the time they are being entered into a study database (standardised template created in Microsoft Excel). At least 10% of data extraction will be double-coded. Discrepancies or difficulties with coding will be resolved through discussion between reviewers or the opinion of a third reviewer will be sought. |
|                      | Non-English-language papers will be excluded (unless data can be obtained from an existing review).   |
|                      | Data Analysis  Where data is available, meta-analysis using a fixed-effects model will be used to combine results from similar studies. Heterogeneity will be considered and if a random-effects model is considered more appropriate it will be conducted.   |

| Topic  | Pharmacological interventions for the treatment of PTSD in adults   |
|--|---|
|  | For risk of bias, outcomes will be downgraded if the randomisation and/or allocation concealment methods are unclear or inadequate. Outcomes will also be downgraded if no attempts are made to blind the assessors or participants in some way, i.e. by either not knowing the aim of the study or the result from other tests. Outcomes will also be downgraded if there is considerable missing data (see below). Handling missing data: |
|  | Where possible an intention to treat approach will be used outcomes will be downgraded if there is a dropout of more than 20%, or if there was a difference of >20% between   |
|  | the groups.  For heterogeneity: outcomes will be downgraded once if I2>50%, twice if I2 >80%  For imprecision: outcomes will be downgraded if:  |
|  | Step 1: If the 95% CI is imprecise i.e. crosses 0.8 or 1.25 (dichotomous) or -0.5 or 0.5 (for continuous). Outcomes will be downgraded one or two levels depending on how many lines it crosses.  |
|  | Step 2: If the clinical decision threshold is not crossed, we will consider whether the criterion for Optimal Information Size is met, if not we will downgrade one level for the following.  |
|  | for dichotomous outcomes: <300 events for continuous outcomes: <400 participants  |
|  | For clinical effectiveness, if studies report outcomes using the same scale mean differences will be considered, if not standardized mean differences (SMDs) will be considered and the following criteria will be used: SMD <0.2 too small to likely show an effect  |
|  | SMD 0.2 small effect SMD 0.5 moderate effect SMD 0.8 large effect   |
|  | RR <0.8 or >1.25 clinical benefit Anything less (RR >0.8 and <1.25), the absolute numbers will be looked at to make a decision on whether there may be a clinical effect.   |
| Heterogeneity (sensitivity analysis and subgroups) | Where substantial heterogeneity exists, sensitivity analyses will be considered, for instance: Studies with <50% completion data (drop out of >50%) will be excluded  |

| Topic | Pharmacological interventions for the treatment of PTSD in adults  |
|-------|--|
|       | Where possible, the influence of subgroups will be considered, including subgroup analyses giving specific consideration to the groups outlined in the sub-question section and to the following groups:  Trauma type (including single incident relative to chronic exposure)  Duration of intervention (for instance, short-term [≤12 weeks] relative to long-term [>12 weeks])  Intensity of intervention (for instance, low intensity [≤15 sessions] relative to high intensity [>15 sessions])  Format of intervention (individual relative to group)  Mode of intervention delivery (including digital relative to face-to-face)  First-line treatment relative to second-line treatment and treatment-resistant PTSD (≥2 inadequate treatments)  Acute PTSD symptoms (clinically important PTSD symptoms for less than 3 months) relative to chronic PTSD symptoms (clinically important PTSD symptoms or more) |
| Notes | Practical and social support (area of scope) is covered quantitatively by interventions listed under psychosocial interventions:  • Supported employment (including individual placement and support [IPS] supported employment and Veterans Health Administration Vocational Rehabilitation Programme [VRP])  • Practical support (including financial and housing)  • Peer support (including self-help groups and support groups)   |

# Appendix B – Literature search strategies

Literature search strategy for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

#### Clinical evidence

Database: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R), Embase, PsycINFO

Date of last search: 29 January 2018

| Date 0 | last search. 29 January 2010   |
|--------|--|
| #      | Searches   |
| 1      | *acute stress/ or *behavioural stress/ or *emotional stress/ or *critical incident stress/ or *mental stress/ or *posttraumatic stress disorder/ or *psychotrauma/   |
| 2      | 1 use emez   |
| 3      | stress disorders, traumatic/ or combat disorders/ or psychological trauma/ or stress disorders, post-traumatic/ or stress disorders, traumatic, acute/ or stress, psychological/   |
| 4      | 3 use mesz, prem   |
| 5      | exp posttraumatic stress disorder/ or acute stress disorder/ or combat experience/ or emotional trauma/ or post-traumatic stress/ or traumatic neurosis/ or trauma/ or psychological stress/ or chronic stress/  |
| 6      | 5 use psyh   |
| 7      | (railway spine or (rape adj2 trauma*) or reexperienc* or re experienc* or torture syndrome or traumatic neuros* or traumatic stress).ti,ab.  |
| 8      | (trauma* and (avoidance or grief or horror or death* or nightmare* or night mare* or emotion*)).ti,ab.   |
| 9      | (posttraumatic* or post traumatic* or stress disorder* or acute stress or ptsd or asd or desnos or (combat neuros* or combat syndrome or concentration camp syndrome or extreme stress or flashback* or flash back* or hypervigilan* or hypervigilen* or psych* stress or psych* trauma* or psycho?trauma* or psychotrauma*) or (posttrauma* or traumagenic* or traumatic stress*)).ti,ab. |
| 10     | or/2,4,6-9   |
| 11     | *psychotherapy/ use emez or psychotherapy/ use mesz, prem,psyh   |
| 12     | (((psycholog* or psycho social* or psychosocial*) adj3 (intervention* or program* or therap* or treat*)) or psychotherap* or psycho therap* or talk* therap* or therapeutic technique* or therapist* or third wave or time limited).ti,ab,sh.  |
| 13     | exp *behavior therapy/ or exp *cognitive therapy/  |
| 14     | 13 use emez  |
| 15     | exp behavior therapy/ use mesz, prem   |
| 16     | exp behavior therapy/ or exp cognitive behavior therapy/   |
| 17     | 16 use psyh  |
| 18     | (((behaviour* or behavior*) adj2 cognitiv*) or cbt or ccbt or ((behav* or cognitive*) adj3 (intervention* or manag* or program* or restructure* or therap* or treat*)) or (stress inoculation adj2 (intervention* or program* or therap* or train* or treat*)) or (behav* adj2   |

| #  | Searches  |
|----|---|
|    | activat*) or ((trauma adj (based or focused or led)) or exposure based or prolonged exposure)).ti,ab.   |
| 19 | *emotion/ use emez or emotions/ use mesz, prem  |
| 20 | emotion focused therapy/ or sympathy/   |
| 21 | 20 use psyh   |
| 22 | (((compassion or emotion* or emotive*) adj (based or focused or led)) or emotional processing or ((compassion or emotion* or emotive*) adj3 (coach* or intervention* or program* or therap* or treat*))).ti,ab.   |
| 23 | exposure therapy/ or narrative therapy/ or virtual reality exposure therapy/  |
| 24 | 23 use emez   |
| 25 | implosive therapy/ or narrative therapy/ or virtual reality exposure therapy/   |
| 26 | 25 use mesz, prem   |
| 27 | exposure therapy/ or narrative therapy/ or virtual reality/   |
| 28 | 27 use psyh   |
| 29 | (((augmented or virtual) adj2 reality) or (virtual adj (environment or restorative)) or ((exposure or implosive or virtual reality) adj2 (intervention* or program* or therap* or train*))).ti,ab.  |
| 30 | ((imagery adj2 (rehears* or re hears*)) or (((lower* or reduc*) adj3 (bad dream* or nightmare*)) and (intervention* or program* or therap* or treat*)) or ((intervention* or program* or therap* or treat*) adj3 nightmare*)).mp. or ((presleep or presleep) adj2 imagery).ti,ab. |
| 31 | (mindfulness or ((exposure or narrative) adj therapy)).sh.  |
| 32 | (kidnet or mindful* or narrative therap*).ti,ab.  |
| 33 | exp "debriefing (psychological)"/ use psyh  |
| 34 | debrief*.ti,ab.   |
| 35 | eye movement desensitization reprocessing/ use mesz, prem or eye movement desensitization therapy/ use psyh or (emdr or (eye movement adj2 desensiti*)).ti,ab.  |
| 36 | hypnosis/ use emez or exp hypnosis/ use mesz, prem or exp hypnotherapy/ use psyh or (hypnosis or hypnotherap*).ti,ab.   |
| 37 | psychodynamic psychotherapy/ use emez or psychotherapy, psychodynamic/ use mesz, prem or psychodynamic psychotherapy/ use psyh or repetitive transcranial magnetic stimulation/ use emez or Transcranial Magnetic Stimulation/ use mesz, prem, psyh                               |
| 38 | ((psychodynamic or (dynamic adj (psychotherapy* or therap*)) or incident reduction) or ((brain or transcranial) adj2 stimulat*) or rtms).ti,ab.   |
| 39 | (psychoanal* or psychosomatic*).ti,ab.  |
| 40 | exp counseling/ use emez,mesz,psyh or counsel*.ti,ab.   |
| 41 | (hg therap* or human givens).ti,ab.   |
| 42 | psychosomatic disorder/th use emez or exp somatoform disorders/th use mesz, prem  |
| 43 | (exp somatoform disorders/ or somatization/) and (intervention* or program* or therap* or treat*).ti,ab,hw. use psyh  |
| 44 | (psychosomatic* or somatherap* or somatic*).ti,ab.  |
| 45 | (emotional freedom or holistic or thought field).ti,ab.   |
| 46 | dance therap*.ti,ab,sh.   |
| 47 | couple therapy/ or family therapy/ or marital therapy/ or exp parent/ed   |
| 48 | 47 use emez   |

| 44 | Convolue  |
|----|---|
| #  | Searches  |
| 49 | couples therapy/ or family therapy/ or marital therapy/ or exp parents/ed   |
| 50 | 49 use mesz, prem   |
| 51 | couples therapy/ or family intervention/ or exp family therapy/ or exp marriage counseling/ or parent training/   |
| 52 | 51 use psyh   |
| 53 | (((con?joint or couple* or family or families or husband* or marriage* or marital* or partner* or relations* or spous* or wife or wives* or (child* adj5 parent*)) adj6 (counsel* or intervention* or program* or support* or therap* or treat*)) or ((couples* or family* or relations*) adj (based or focused or led)) or ecological therap* or expressed emotion or family dynamics or family relationships).tw.   |
| 54 | ((child* adj2 family traumatic stress intervention) or cftsi).ti,ab.  |
| 55 | play therapy.sh.  |
| 56 | (doll therap* or ((play or playful) adj3 (intervention* or program* or therap* or treat*)) or sandplay or sand play).ti,ab.   |
| 57 | meditation.sh. or meditat*.ti,ab.   |
| 58 | mindfulness.sh. or (mbsr or mindful*).ti,ab.  |
| 59 | exp horticulture/ or occupational therapy/ or recreational therapy/   |
| 60 | 59 use emez   |
| 61 | horticultural therapy/ or occupational therapy/ or recreation therapy/  |
| 62 | 61 use mesz, prem   |
| 63 | exp "nature (environment)"/ or horticulture therapy/ or recreation therapy/ or occupational therapy/  |
| 64 | 63 use psyh   |
| 65 | ((nature adj (assisted or based)) or (nature adj3 (intervention* or program* or therap* or treat*)) or ecotherap* or e cotherap* or gardening or horticult* or leisure activit* or naturopath* or occupational therap*).ti,ab. or exp animal assisted therapy/ use emez, mesz or animal assisted therapy/ use psyh or (((animal* or dog* or equine* or horse* or pet or pets) adj2 (assist* or based or facilitat*)) or ((animal* or dog* or equine* or horse* or pet or pets) adj3 (intervention* or therap* or treat* or program*))).ti,ab. |
| 66 | psychoeducation.sh. or (psychoed* or psycho ed*).ti,ab.   |
| 67 | exp acupuncture/ use emez or exp alternative medicine/ use emez or biofeedback/ or massage/ use emez or meditation/ use emez or acupressure/ use mesz, prem or massage/ use mesz, prem or acupuncture/ use mesz, prem or exp complementary therapies/ use mesz, prem or exp alternative medicine/ use psyh or biofeedback/ use psyh or massage/ use psyh or mind body therapy/ use psyh   |
| 68 | (chinese medicine or medicine, chinese traditional or (moxibustion or electroacupuncture)).sh,id. or ((alternative or complementary) adj2 (medicine* or therap*)).ti,ab,sh. or (acu point* or acupoint* or acupressur* or acupunctur* or (ching adj2 lo) or cizhen or dianzhen or electroacupunctur* or (jing adj2 luo) or jingluo or massag* or needle therap* or tapping or zhenjiu or zhenci).tw.  |
| 69 | exp *exercise/ use emez or exp *kinesiotherapy/ use emez or exp exercise/ use mesz, prem or exercise therapy/ use mesz, prem or exp exercise/ use psyh (physiotherap* or physiotherap* or rehab*).ti,ab,hw.   |
| 70 | (((balance or flexibility or resistance or sitting* or strenth*) adj2 (exercise* or train*)) or aerobic* or anaerobic* or bowls or dancing or dance or cycling or cycle* or elliptical train* or jogging or low impact activit* or running or swimming or sprinting or swim*1 or walking or   |

| #  | Searches   |
|----|--|
|    | yoga or tai chi or weight train* or (weight and brain* and (change* or increas* or volum*))).ti,ab.  |
| 71 | friendship/ or peer counseling/ or peer group/ or self help/ or self care/ or social network/ or social support/ or support group/   |
| 72 | 71 use emez  |
| 73 | community networks/ or friends/ or exp peer group/ or self care/ or self-help groups/ or social networking/ or social support/   |
| 74 | 73 use mesz, prem  |
| 75 | friendship/ or network therapy/ or exp social networks/ or peer relations/ or peers/ or peer counseling/ or self care skills/ or exp self help techniques/ or social support/ or exp support groups/   |
| 76 | 75 use psyh  |
| 77 | ((self adj (administer* or assess* or attribut* or care or change or directed or efficacy or help* or guide* or instruct* or manag* or medicat* or monitor* or regulat* or reinforc* or re inforc* or support* or technique* or therap* or train* or treat*)) or selfadminister* or selfassess* or selfattribut* or selfcare or selfchange or selfdirected or selfefficacy or selfhelp* or selfguide* or selfinstruct* or selfmanag* or selfmedicat* or selfmonitor* or selfregulat* or selfreinforc* or self re inforc* or selfsupport* or selftechnique* or selftherap* or selftrain* or selftreat* or (wellness adj (therap* or train* or treat*))).ti,ab,sh.   |
| 78 | (befriend* or be*1 friend* or buddy or buddies or ((community or lay or paid or support) adj (person or worker*))).ti,ab.  |
| 79 | (((consumer* or famil* or friend* or lay or mutual* or peer* or social* or spous* or voluntary or volunteer*) adj3 (assist* or advice* or advis* or counsel* or educat* or forum* or help* or mentor* or network* or support* or visit*)) or ((consumer* or famil* or peer* or self help or social* or support* or voluntary or volunteer*) adj2 group*) or ((consumer* or famil* or friend* or lay or mutual* or peer* or self help or social* or spous* or support* or voluntary or volunteer*) adj3 (intervention* or program* or rehab* or therap* or service* or skill* or treat*)) or (((consumer* or famil* or friend* or lay* or peer* or spous* or user* or support* or voluntary or volunteer*) adj (based or counsel* or deliver* or interact* or led or mediat* or operated or provides or provider* or run*)) or ((consumer* or famil* or friend* or lay* or peer* or relation* or spous* or support*) adj3 trust*) or voluntary work*)).ti,ab. |
| 80 | (((lay or peer*) adj3 (advis* or consultant or educator* or expert* or facilitator* or instructor* or leader* or mentor* or person* or tutor* or worker*)) or expert patient* or mutual aid).ti,ab.  |
| 81 | (peer* adj3 (assist* or counsel* or educat* or program* or rehab* or service* or supervis*)).ti,ab.  |
| 82 | ((psychoeducat* or psycho educat*) adj3 (group or network* or service*)).ti,ab.  |
| 83 | ((psychosocial or social) adj work*).ti,ab.  |
| 84 | ((ptsd or posttrauma* or post trauma* or trauma*) adj2 support*).ti,ab.  |
| 85 | recovery support.ti,ab.  |
| 86 | financial management/ use emez or financial support/ use mesz, prem or finance/ use psyh   |
| 87 | ((financ* or money) adj2 (assist* or educat* or guidance or intervention* or program* or support* or train*)).ti,ab.   |
| 88 | assisted living facility/ or emergency shelter/ or halfway house/ or housing/ or independent living/ or residential home/ or residential home/   |
| 89 | 88 use emez  |
| 90 | assisted living facilities/ or emergency shelter/ or group homes/ or halfway houses/ or housing/ or independent living/ or residential facilities/   |

| #   | Searches  |
|-----|---|
| 91  | 90 use mesz, prem   |
| 92  | assisted living / use psyh or shelters/ use psyh or group homes/ use psyh or halfway houses/ use psyh or housing/ use psyh or residential care institutions/ use psyh or ((resident* or hous* or accommod* or commun* or comu* or home*) adj5 (support* or support* or shelter* or outreach* or visit* or appointment*)).ti,ab. |
| 93  | (residential treatm* or residential facility* or supported hous* or public hous*).ti,ab.  |
| 94  | (accomod* or assertive community treatment* or home* or housing* or outreach* or residential*).ti,ab.   |
| 95  | absenteeism/ or daily life activity/ or employment/ or medical leave/ or mentoring/ or occupational health/ or occupational therapy/ or return to work/ or supported employment/ or unemployment/ or vocational guidance/ or vocational rehabilitation/ or work capacity/ or work/  |
| 96  | 95 use emez   |
| 97  | absenteeism/ or "activities of daily living"/ or employment, supported/ or employment/ or mentoring/ or occupational health/ or occupational therapy/ or rehabilitation, vocational/ or return to work/ or sick leave/ or unemployment/ or vocational guidance/ or work/  |
| 98  | 97 use mesz, prem   |
| 99  | "activities of daily living"/ or exp coaching/ or employee absenteeism/ or employment status/ or occupational guidance/ or occupational health/ or occupational therapy/ or reemployment/ or unemployment/ or vocational counselors/ or exp vocational rehabilitation/  |
| 100 | 99 use psyh   |
| 101 | (((supp* or transitional*) adj5 (employ* or work*)) or individual placement or (placement* adj3 (employ* or work*))).ti,ab.   |
| 102 | ((employ* or placement* or psychosocial* or psycho-social* or occupation* or soc* or vocation* or work* or job* or counsel*) adj5 rehab*).ti,ab.  |
| 103 | (sheltered work* or vocatio* or fountain house* or fountainhouse* or clubhouse* or clubhouse* or work therap*).ti,ab.   |
| 104 | (transitional employment or rehabilitation counsel* or (occupational adj (health or medicine)) or work* adjustment).ti,ab.  |
| 105 | ((performance adj (activit* or coach* or management or occupation*)) or coaching).ti,ab.  |
| 106 | (((sheltered or permitted or voluntary or vocational or return* or rehabilitat*) adj3 work*) or work capacity or reemploy* or re employ* or job retention or work capacity).ti,ab.  |
| 107 | ((employ* or job or occupation* or vocation* or work*) adj5 (counsel* or educat* or guidance* or intervention* or program* or rehab* or reintegrat* or re integrat* or support* or therap* or train*)).ti,ab.   |
| 108 | placement.ti,ab.  |
| 109 | or/11-12,14-15,17-19,21-22,24,26,28-46,48,50,52-58,60,62,64-70,72,74,76-87,89,91-94,96,98,100-108   |
| 110 | meta analysis/ or "meta analysis (topic)"/ or systematic review/  |
| 111 | 110 use emez  |
| 112 | meta analysis.sh,pt. or "meta-analysis as topic"/ or "review literature as topic"/  |
| 113 | 112 use mesz, prem  |
| 114 | (literature review or meta analysis).sh,id,md. or systematic review.id,md.  |
| 115 | 114 use psyh  |
| 116 | (exp bibliographic database/ or (((electronic or computer* or online) adj database*) or bids or cochrane or embase or index medicus or isi citation or medline or psyclit or psychlit or  |

| #   | Searches   |
|-----|--|
|     | scisearch or science citation or (web adj2 science)).ti,ab.) and (review*.ti,ab,sh,pt. or  |
|     | systematic*.ti,ab.)  |
| 117 | 116 use emez   |
| 118 | (exp databases, bibliographic/ or (((electronic or computer* or online) adj database*) or bids or cochrane or embase or index medicus or isi citation or medline or psyclit or psychlit or scisearch or science citation or (web adj2 science)).ti,ab.) and (review*.ti,ab,sh,pt. or systematic*.ti,ab.) |
| 119 | 118 use mesz, prem   |
| 120 | (computer searching.sh,id. or (((electronic or computer* or online) adj database*) or bids or cochrane or embase or index medicus or isi citation or medline or psyclit or psychlit or scisearch or science citation or (web adj2 science)).ti,ab.) and (review*.ti,ab,pt. or systematic*.ti,ab.)        |
| 121 | 120 use psyh   |
| 122 | ((analy* or assessment* or evidence* or methodol* or quantativ* or systematic*) adj2 (overview* or review*)).tw. or ((analy* or assessment* or evidence* or methodol* or quantativ* or systematic*).ti. and review*.ti,pt.) or (systematic* adj2 search*).ti,ab.   |
| 123 | (metaanal* or meta anal*).ti,ab.   |
| 124 | (research adj (review* or integration)).ti,ab.   |
| 125 | reference list*.ab.  |
| 126 | bibliograph*.ab.   |
| 127 | published studies.ab.  |
| 128 | relevant journals.ab.  |
| 129 | selection criteria.ab.   |
| 130 | (data adj (extraction or synthesis)).ab.   |
| 131 | (handsearch* or ((hand or manual) adj search*)).ti,ab.   |
| 132 | (mantel haenszel or peto or dersimonian or der simonian).ti,ab.  |
| 133 | (fixed effect* or random effect*).ti,ab.   |
| 134 | ((pool* or combined or combining) adj2 (data or trials or studies or results)).ti,ab.  |
| 135 | or/111,113,115,117,119,121-134   |
| 136 | exp "clinical trial (topic)"/ or exp clinical trial/ or crossover procedure/ or double blind procedure/ or placebo/ or randomization/ or random sample/ or single blind procedure/   |
| 137 | 136 use emez   |
| 138 | exp clinical trial/ or exp "clinical trials as topic"/ or cross-over studies/ or double-blind method/ or placebos/ or random allocation/ or single-blind method/   |
| 139 | 138 use mesz, prem   |
| 140 | (clinical trials or placebo or random sampling).sh,id.   |
| 141 | 140 use psyh   |
| 142 | (clinical adj2 trial*).ti,ab.  |
| 143 | (crossover or cross over).ti,ab.   |
| 144 | (((single* or doubl* or trebl* or tripl*) adj2 blind*) or mask* or dummy or doubleblind* or singleblind* or trebleblind* or tripleblind*).ti,ab.   |
| 145 | (placebo* or random*).ti,ab.   |
| 146 | treatment outcome*.md. use psyh  |
| 147 | animals/ not human*.mp. use emez   |

| #   | Searches                            |
|-----|-------------------------------------|
| 148 | animal*/ not human*/ use mesz, prem |
| 149 | (animal not human).po. use psyh     |
| 150 | or/137,139,141-146                  |
| 151 | 150 not (or/147-149)                |
| 152 | or/135,151                          |
| 153 | 10 and 109 and 152                  |
|     |                                     |

Database: CDSR, DARE, HTA, CENTRAL

Date of last search: 29 January 2018

| #   | Searches   |
|-----|--|
| #1  | MeSH descriptor: Stress Disorders, Traumatic this term only  |
| #2  | MeSH descriptor: Combat Disorders this term only   |
| #3  | MeSH descriptor: Psychological Trauma this term only   |
| #4  | MeSH descriptor: Stress Disorders, Post-Traumatic this term only   |
| #5  | MeSH descriptor: Stress Disorders, Traumatic, Acute this term only   |
| #6  | MeSH descriptor: Stress, Psychological this term only  |
| #7  | ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress"):ti (Word variations have been searched)   |
| #8  | ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress"):ab (Word variations have been searched)   |
| #9  | (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)):ti (Word variations have been searched)  |
| #10 | (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)):ab (Word variations have been searched)  |
| #11 | (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttrauma* or traumagenic* or "traumatic stress*")):ti (Word variations have been searched) |
| #12 | (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttrauma* or traumagenic* or "traumatic stress*")):ab (Word variations have been searched) |
| #13 | #1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12  |
|     |  |

Database: CINAHL PLUS

Date of last search: 29 January 2018

| #   | Searches    |
|-----|-------------|
| s52 | s6 and s51  |
| s51 | s40 or s50  |
| s50 | s48 not s49 |

| #   | Searches   |
|-----|--|
| s49 | (mh "animals") not (mh "human")  |
| s48 | s41 or s42 or s43 or s44 or s45 or s46 or s47  |
| s47 | ti ( placebo* or random* ) or ab ( placebo* or random* )   |
| s46 | ti ( single blind* or double blind* or treble blind* or mask* or dummy* or singleblind* or doubleblind* or trebleblind* ) or ab ( single blind* or double blind* or treble blind* or mask* or dummy* or singleblind* or doubleblind* or trebleblind* or tripleblind* )   |
| s45 | ti ( crossover or cross over ) or ab ( crossover or cross over )   |
| s44 | ti clinical n2 trial* or ab clinical n2 trial*   |
| s43 | (mh "crossover design") or (mh "placebos") or (mh "random assignment") or (mh "random sample")   |
| s42 | mw double blind* or single blind* or triple blind*   |
| s41 | (mh "clinical trials+")  |
| s40 | s7 or s8 or s9 or s10 or s11 or s12 or s13 or s14 or s15 or s16 or s17 or s18 or s19 or s20 or s21 or s22 or s23 or s29 or s30 or s31 or s34 or s35 or s36 or s37 or s38 or s39  |
| s39 | ti ( analy* n5 review* or evidence* n5 review* or methodol* n5 review* or quantativ* n5 review* or systematic* n5 review* ) or ab ( analy* n5 review* or assessment* n5 review* or evidence* n5 review* or methodol* n5 review* or qualitativ* n5 review* or quantativ* n5 review* or systematic* n5 review* ) |
| s38 | ti ( pool* n2 results or combined n2 results or combining n2 results ) or ab ( pool* n2 results or combined n2 results or combining n2 results )   |
| s37 | ti ( pool* n2 studies or combined n2 studies or combining n2 studies ) or ab ( pool* n2 studies or combined n2 studies or combining n2 studies )   |
| s36 | ti ( pool* n2 trials or combined n2 trials or combining n2 trials ) or ab ( pool* n2 trials or combined n2 trials or combining n2 trials )   |
| s35 | ti ( pool* n2 data or combined n2 data or combining n2 data ) or ab ( pool* n2 data or combined n2 data or combining n2 data )   |
| s34 | s32 and s33  |
| s33 | ti review* or pt review*   |
| s32 | ti analy* or assessment* or evidence* or methodol* or quantativ* or qualitativ* or systematic*   |
| s31 | ti "systematic* n5 search*" or ab "systematic* n5 search*"   |
| s30 | ti "systematic* n5 review*" or ab "systematic* n5 review*"   |
| s29 | (s24 or s25 or s26) and (s27 or s28)   |
| s28 | ti systematic* or ab systematic*   |
| s27 | tx review* or mw review* or pt review*   |
| s26 | (mh "cochrane library")  |
| s25 | ti ( bids or cochrane or embase or "index medicus" or "isi citation" or medline or psyclit or psychlit or scisearch or "science citation" or web n2 science ) or ab ( bids or cochrane or "index medicus" or "isi citation" or psyclit or psychlit or scisearch or "science citation" or web n2 science )      |
| s24 | ti ("electronic database*" or "bibliographic database*" or "computeri?ed database*" or "online database*") or ab ("electronic database*" or "bibliographic database*" or "computeri?ed database*" or "online database*")   |
| s23 | (mh "literature review")   |
| s22 | pt systematic* or pt meta*   |
| s21 | ti ( "fixed effect*" or "random effect*" ) or ab ( "fixed effect*" or "random effect*" )   |

| #   | Convolue  |
|-----|---|
| #   | Searches ""   |
| s20 | ti ( "mantel haenszel" or peto or dersimonian or "der simonian" ) or ab ( "mantel haenszel" or peto or dersimonian or "der simonian" )  |
| s19 | ti ( handsearch* or "hand search*" or "manual search*" ) or ab ( handsearch* or "hand search*" )  |
| s18 | ab "data extraction" or "data synthesis"  |
| s17 | ab "selection criteria"   |
| s16 | ab "relevant journals"  |
| s15 | ab "published studies"  |
| s14 | ab bibliograph*   |
| s13 | ti "reference list*"  |
| s12 | ab "reference list*"  |
| s11 | ti ( "research review*" or "research integration" ) or ab ( "research review*" or "research integration" )  |
| s10 | ti ( metaanal* or "meta anal*" or metasynthes* or "meta synethes*" ) or ab ( metaanal* or "meta anal*" or metasynthes* or "meta synethes*" )  |
| s9  | (mh "meta analysis")  |
| s8  | (mh "systematic review")  |
| s7  | (mh "literature searching+")  |
| s6  | s1 or s2 or s3 or s4 or s5  |
| s5  | ti ( (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttraumatic* or "traumatic stress*")) ) or ab ( (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttrauma* or traumagenic* or "traumatic stress*")) ) |
| s4  | ti ( (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)) ) or ab ( (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)) )  |
| s3  | ti ( ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress") ) or ab ( ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress") )  |
| s2  | (mh "stress, psychological")  |
| s1  | (mh "stress disorders, post-traumatic")   |
|     |   |

#### Health economic evidence

Note: evidence resulting from the health economic search update was screened to reflect the final dates of the searches that were undertaken for the clinical reviews (see review protocols).

Database: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R), Embase, PsycINFO

#### Date of last search: 1 March 2018

|    | i last search. I March 2016  |
|----|--|
| #  | Searches   |
| 1  | *acute stress/ or *behavioural stress/ or *emotional stress/ or *critical incident stress/ or *mental stress/ or *posttraumatic stress disorder/ or *psychotrauma/   |
| 1  | *acute stress/ or *behavioural stress/ or *emotional stress/ or *critical incident stress/ or *mental stress/ or *posttraumatic stress disorder/ or *psychotrauma/   |
| 2  | 1 use emez   |
| 3  | stress disorders, traumatic/ or combat disorders/ or psychological trauma/ or stress disorders, post-traumatic/ or stress disorders, traumatic, acute/ or stress, psychological/   |
| 4  | 3 use mesz, prem   |
| 5  | exp posttraumatic stress disorder/ or acute stress disorder/ or combat experience/ or "debriefing (psychological)"/ or emotional trauma/ or post-traumatic stress/ or traumatic neurosis/ or "trauma"/ or stress reactions/ or psychological stress/ or chronic stress/  |
| 6  | 5 use psyh   |
| 7  | (railway spine or (rape adj2 trauma*) or reexperienc* or re experienc* or torture syndrome or traumatic neuros* or traumatic stress).ti,ab.  |
| 8  | (trauma* and (avoidance or grief or horror or death* or nightmare* or night mare* or emotion*)).ti,ab.   |
| 9  | (posttraumatic* or post traumatic* or stress disorder* or acute stress or ptsd or asd or desnos or (combat neuros* or combat syndrome or concentration camp syndrome or extreme stress or flashback* or flash back* or hypervigilan* or hypervigilen* or psych* stress or psych* trauma* or psycho?trauma* or psychotrauma*)).ti,ab. |
| 10 | or/2,4,6-9   |
| 11 | budget/ or exp economic evaluation/ or exp fee/ or funding/ or exp health care cost/ or health economics/ or exp pharmacoeconomics/ or resource allocation/  |
| 12 | 151 use emez   |
| 13 | exp budgets/ or exp "costs and cost analysis"/ or economics/ or exp economics, hospital/ or exp economics, medical/ or economics, nursing/ or economics, pharmaceutical/ or exp "fees and charges"/ or value of life/  |
| 14 | 153 use mesz, prem   |
| 15 | exp "costs and cost analysis"/ or cost containment/ or economics/ or finance/ or funding/ or "health care economics"/ or pharmacoeconomics/ or exp professional fees/ or resource allocation/  |
| 16 | 155 use psyh   |
| 17 | (cost* or economic* or pharmacoeconomic* or pharmaco economic*).ti. or (cost* adj2 (effective* or utilit* or benefit* or minimi*)).ab. or (budget* or fee or fees or financ* or price or prices or pricing or resource* allocat* or (value adj2 (monetary or money))).ti,ab.   |
| 18 | or/12,14,16-17   |
| 19 | decision theory/ or decision tree/ or monte carlo method/ or nonbiological model/ or (statistical model/ and exp economic aspect/) or stochastic model/ or theoretical model/  |
| 20 | 159 use emez   |
| 21 | exp decision theory/ or markov chains/ or exp models, economic/ or models, organizational/ or models, theoretical/ or monte carlo method/  |
| 22 | 161 use mesz, prem   |
| 23 | exp decision theory/ or exp stochastic modeling/   |
| 24 | 163 use psyh   |
| 25 | ((decision adj (analy* or model* or tree*)) or economic model* or markov).ti,ab.   |
|    |  |

| #  | Searches   |
|----|--|
| 26 | or/20,22,24-25   |
| 27 | quality adjusted life year/ or "quality of life index"/ or short form 12/ or short form 20/ or short form 36/ or short form 8/ or sickness impact profile/ |
| 28 | 167 use emez   |
| 29 | quality-adjusted life years/ or sickness impact profile/   |
| 30 | 169 use mesz, prem   |
| 31 | (((disability or quality) adj adjusted) or (adjusted adj2 life)).ti,ab.  |
| 32 | (disutili* or dis utili* or (utilit* adj1 (health or score* or value* or weigh*))).ti,ab.  |
| 33 | (health year equivalent* or hye or hyes).ti,ab.  |
| 34 | (daly or qal or qale or qaly or qtime* or qwb*).ti,ab.   |
| 35 | discrete choice.ti,ab.   |
| 36 | (euroqol* or euro qol* or eq5d* or eq 5d*).ti,ab.  |
| 37 | (hui or hui1 or hui2 or hui3).ti,ab.   |
| 38 | (((general or quality) adj2 (wellbeing or well being)) or quality adjusted life or qwb or (value adj2 (money or monetary))).ti,ab.                         |
| 39 | (qol or hql* or hqol* or hrql or hrql).ti,ab.  |
| 40 | rosser.ti,ab.  |
| 41 | sickness impact profile.ti,ab.   |
| 42 | (standard gamble or time trade* or tto or willingness to pay or wtp).ti,ab.  |
| 43 | (sf36 or sf 36 or short form 36 or shortform 36 or shortform36).ti,ab.   |
| 44 | (sf6 or sf 6 or short form 6 or shortform 6 or shortform6).ti,ab.  |
| 45 | (sf12 or sf 12 or short form 12 or shortform 12 or shortform12).ti,ab.   |
| 46 | (sf16 or sf 16 or short form 16 or shortform 16 or shortform16).ti,ab.   |
| 47 | (sf20 or sf 20 or short form 20 or shortform 20 or shortform20).ti,ab.   |
| 48 | (sf8 or sf 8 or short form 8 or shortform 8 or shortform8).ti,ab.  |
| 49 | or/28,30-48  |
| 50 | or/18,26,49  |

### Database: HTA, NHS EED

Date of last search: 1 March 2018

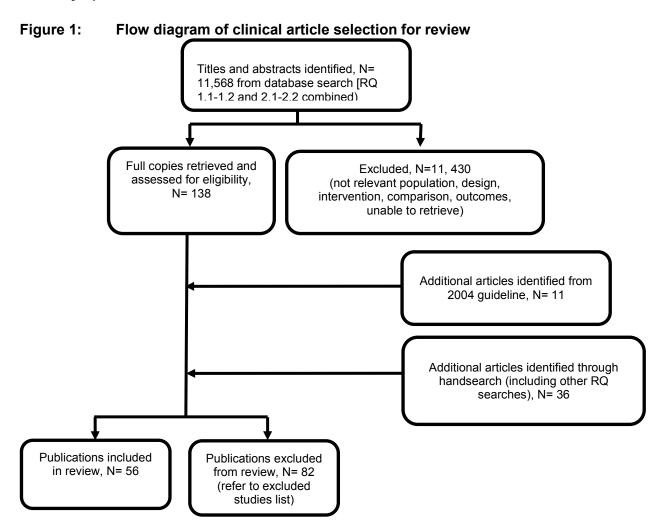
| _ | alc 0 | That Scaron. I Waren 2010  |
|---|-------|--|
|   | #     | Searches   |
|   | #1    | MeSH descriptor: Stress Disorders, Traumatic this term only  |
|   | #2    | MeSH descriptor: Combat Disorders this term only   |
|   | #3    | MeSH descriptor: Psychological Trauma this term only   |
|   | #4    | MeSH descriptor: Stress Disorders, Post-Traumatic this term only   |
|   | #5    | MeSH descriptor: Stress Disorders, Traumatic, Acute this term only   |
|   | #6    | MeSH descriptor: Stress, Psychological this term only  |
|   | #7    | ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress"):ti (Word variations have been searched) |

| #   | Searches   |
|-----|--|
| #8  | ("railway spine" or (rape near/2 trauma*) or reexperienc* or "re experienc*" or "torture syndrome" or "traumatic neuros*" or "traumatic stress"):ab (Word variations have been searched)   |
| #9  | (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)):ti (Word variations have been searched)  |
| #10 | (trauma* and (avoidance or grief or horror or death* or nightmare* or "night mare*" or emotion*)):ab (Word variations have been searched)  |
| #11 | (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttrauma* or traumagenic* or "traumatic stress*")):ti (Word variations have been searched) |
| #12 | (posttraumatic* or "post traumatic*" or "stress disorder*" or "acute stress" or ptsd or asd or desnos or ("combat neuros*" or "combat syndrome" or "concentration camp syndrome" or "extreme stress" or flashback* or "flash back*" or hypervigilan* or hypervigilen* or "psych* stress" or "psych* trauma*" or psychotrauma* or psychotrauma*) or (posttrauma* or traumagenic* or "traumatic stress*")):ab (Word variations have been searched) |
| #13 | #1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12  |



# **Appendix C – Clinical evidence study selection**

Clinical evidence study selection for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"



# **Appendix D – Clinical evidence tables**

Clinical evidence tables for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

## **Psychological: Trauma-focused CBT**

| Study ID    | Intervention  | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria  |
|-------------|---|---|--|----|--|---|
| Ahrens 2002 | Trauma-focused CBT: Cohen TF-CBT/Cognitive processing therapy | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Mixed - Adolescent offenders incarcerated in a youth facility. Interview data indicated that about one-third of the youths had experienced multiple traumas (n = 11, 29%), and over half had documented trauma histories (n = 26 or 68%, as documented in their charts from collateral sources ranging from Social Rehabilitation Service investigations, child protective services reports, hospital reports, etc.) | 38 | Age range (mean): 15-18 (16.4) Gender (% female): 0 BME (% non-white): 40 Country: US Coexisting conditions: 52% stated they had experienced a head injury that led to loss of consciousness; 40% stated that they had been diagnosed with ADD or ADHD in the past Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): 29% had experienced multiple traumas; 68% had documented trauma histories (in their charts | Inclusion criteria: Adolescent males incarcerated in a youth facility for adolescent offenders who met DSM-IV criteria for PTSD |

| Study ID        | Intervention   | PTSD details  | Trauma type   | N  | Demographics   | Inclusion/Exclusion criteria   |
|-----------------|--|---|---|----|--|--|
|                 |  |   |   |    | from collateral sources ranging from Social Rehabilitation Service investigations, child protective services reports, hospital reports, etc.). Single or multiple incident index trauma: Multiple  |  |
| Al-Hadethe 2015 | Trauma-focused<br>CBT: Narrative<br>exposure therapy<br>for traumatized<br>children and<br>adolescents<br>(KidNET) | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Unclear (Not reported in details)   | 60 | Age range (mean): 16-19 (NR) Gender (% female): 0 BME (% non-white): Unclear Country: Iraq Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Unclear | Inclusion criteria: male secondary school students aged 16-19 years old, who met DSM-IV criteria for PTSD as measured by the Scale of Posttraumatic Stress Symptoms (SPTSS)  |
| Auslander 2017  | Trauma-focused<br>CBT: CBT group   | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Mixed (Girls involved in<br>child welfare who had<br>histories of abuse and<br>neglect. Girls with<br>histories of sexual<br>abuse were included) | 34 | Age range (mean): 12-18 (14.6) Gender (% female): 100 BME (% non-white): 78 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR  | Inclusion criteria: girls who had histories of child maltreatment investigated by child protective services; aged 12-18 years; reported histories of trauma with corresponding symptoms that were causing emotional, psychological, and/or relationship difficulties (based on the observations and assessments of their referring caseworker or therapist). Exclusion |

| Study ID                         | Intervention  | PTSD details  | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria  |
|----------------------------------|---|---|--|-----|---|---|
|                                  |   |   |  |     | Single or multiple incident index trauma: Multiple  | criteria: severe learning problems (i.e., could not read or write), active suicidal or psychotic thoughts, or had severe behavioural disorders that would prohibit their participation in a group or interview. Participants who were recently hospitalized for mental health problems were delayed entry into the study (after a 6-month waiting period) |
| Berger 2009                      | Trauma-focused<br>CBT: CBT group  | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Natural disasters (such as severe floods, earthquakes or tsunamis) - Tsunami (Sri Lanka, December 26 2004) - 84% present and physically hurt during the tsunami; 12% present during the tsunami, but were not hurt; 4% not personally exposed to the tsunami. 89.2% had been exposed to a major traumatic incident not related to the tsunami. | 166 | Age range (mean): 9-14 (NR) Gender (% female): 48 BME (% non-white): NR Country: Sri Lanka Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion: children aged 9-14 years exposed to the 2004 tsunami and DSMIVTR (One positive response regarded as meeting criterion A1 of PTSD, and one score of at least 4 was necessary to fulfil criterion A2 of PTSD)  |
| Catani<br>2009/Rockstroh<br>2004 | Trauma-focused CBT: Narrative exposure therapy for traumatized children and | Clinically important PTSD symptoms (scoring above                                 | Natural disasters (such<br>as severe floods,<br>earthquakes or<br>tsunamis) - Tsunami<br>disaster in Sri Lanka   | 31  | Age range (mean): 8-<br>14(11.9)<br>Gender (% female): 45<br>BME (% non-white): NR  | Inclusion criteria: children within refugee camps following tsunami who met criteria for a preliminary PTSD diagnosis (all DSM-IV criteria except time criterion).  |

| Study ID            | Intervention  | PTSD details   | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria  |
|---------------------|---|--|--|----|--|---|
|                     | adolescents<br>(KidNET)   | a threshold on validated scale)  |  |    | Country: Sri Lanka Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean number of traumas 4.6. 81% identified the tsunami as the worst traumatic event experienced but 68% had also been affected by traumatic war experiences Single or multiple incident index trauma: Multiple | Exclusion criteria: mental retardation, psychosis or any neurological disorder  |
| Chen 2014           | Trauma-focused<br>CBT: CBT group  | Clinically<br>important PTSD<br>symptoms<br>(scoring above<br>a threshold on<br>validated scale) | Natural disasters (such<br>as severe floods,<br>earthquakes or<br>tsunamis): Adolescents<br>who had lost at least 1<br>parent in the Sichuan,<br>China, Earthquake | 40 | Age range (mean): NR (14.5) Gender (% female): 68 BME (% non-white): NR Country: Israel Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single   | Inclusion criteria: adolescents from<br>two secondary schools, who had<br>lost at least one parent in the<br>earthquake, and scored≥18 on the<br>CRIES-13       |
| Cohen<br>1998/2005a | Trauma-focused<br>CBT: Cohen TF-<br>CBT/Cognitive<br>processing therapy | Clinically<br>important PTSD<br>symptoms<br>(scoring above                                       | Childhood sexual<br>abuse - Contact sexual<br>abuse perpetrated by<br>someone at least 5<br>years older than the   | 82 | Age range (mean): 7-<br>15(11.1)<br>Gender (% female): 69<br>BME (% non-white): 41   | Inclusion criteria: contact sexual abuse within the past 6 months which had been validated by CPS or an independent forensic evaluation prior to entry into the |

| Study ID                         | Intervention  | PTSD details  | Trauma type   | N   | Demographics   | Inclusion/Exclusion criteria   |
|----------------------------------|---|---|---|-----|--|--|
|                                  |   | a threshold on validated scale)   | participants (36% single episode, 21% 2-5 abuse occasions, 8% 6-10 times, 33% were abused more than 10 times; 2% unknown) |     | Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple   | study, significant symptomatology related to the sexual abuse experience (as documented by a score in the clinical range on any of the self-report instruments or the presence of sexually inappropriate behaviour as reported by the parent), and availability of a non-offending parent or primary caregiver who was able to participate in treatment. Exclusion criteria: Active psychotic symptoms or substance abuse, or mental retardation or pervasive developmental delay in the child, or active psychosis in the parent or primary caretaker participating in the treatment. |
| Cohen<br>2004a/Deblinger<br>2006 | Trauma-focused<br>CBT: Narrative<br>exposure therapy<br>(NET) | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Childhood sexual<br>abuse (Contact sexual<br>abuse)   | 229 | Age range (mean): 8- 14(10.8) Gender (% female): 79 BME (% non-white): 40 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean 2.66 (SD 1.61) traumatic events in addition to sexual abuse. Previous trauma: 58% had witnessed domestic | Inclusion criteria: children who had experienced contact sexual abuse that was confirmed by Child Protective Services (CPS), law enforcement, or a professional independent forensic evaluator; who met at least five criteria for sexual abuse—related DSM-IV-defined PTSD, including at least one symptom in each of the three PTSD clusters (re-experiencing, avoidance or numbing, and hyperarousal); who had a parent or other caretaker (including long-term foster parents) who was willing and able to participate in the  |

| Study ID            | Intervention  | PTSD details   | Trauma type  | N   | Demographics   | Inclusion/Exclusion criteria  |
|---------------------|---|--|--|-----|--|---|
|                     |   |  |  |     | violence, 26% were victims of physical abuse, 37% had witnessed or been involved in a serious accident, 17% were victims or witnesses of community violence, 14% had experienced a fire or natural disaster, and 25% had experienced other PTSD-level traumatic events, such as medical traumas, traumatic custody situations (e.g., being kidnapped by noncustodial parent), school violence not included in the K-SADS definition of community violence, and terrorist attacks  Single or multiple incident index trauma: Multiple | parental treatment component of the study. Exclusion criteria: an active psychotic disorder or an active substance use disorder that resulted in significant impairment in adaptive functioning, or if the parent or primary caretaker who would be participating in the treatment had such a disorder; non-fluency in English; a documented developmental disorder (e.g., autism); children who were currently taking psychotropic medication who had not been on a stable medication regimen for at least 2 months prior to admission to the study. |
| Cohen<br>2011/2005b | Trauma-focused<br>CBT: Cohen TF-<br>CBT/Cognitive<br>processing therapy | Clinically<br>important PTSD<br>symptoms<br>(scoring above<br>a threshold on<br>validated scale) | Domestic violence<br>(Children exposed to<br>intimate partner<br>violence) | 124 | Age range (mean): 7-14 (9.6) Gender (% female): 51 BME (% non-white): 44 Country: US Coexisting conditions: Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean  | Inclusion criteria: children aged 7-<br>14 years; had at least 5 IPV-<br>related PTSD symptoms, including<br>at least 1 in each of 3 PTSD<br>symptom clusters on the Kiddie<br>Schedule for Affective Disorders<br>and Schizophrenia, Present and<br>Lifetime Version (K-SADS-PL);<br>were fluent in English and had an<br>English-speaking mother who was  |

| Study ID               | Intervention  | PTSD details  | Trauma type  | N   | Demographics   | Inclusion/Exclusion criteria   |
|------------------------|---|---|--|-----|--|--|
|                        |   |   |  |     | number of trauma types: 3.7. Past trauma experiences: Car accident (15%); Other accident (38%); Fire (12%); Disaster (9%); Witness to violent crime (23%); Victim of violent crime (18%); Physical abuse (36%); Sexual abuse (8%); Other (44%) Single or multiple incident index trauma: Multiple  | a direct IPV victim; assented (and their mother consented) to participate in 8 therapy sessions. Exclusion criteria: a significant developmental disorder or an IQ less than 80; serious psychotic symptoms in parent or child; living in an IPV shelter.  |
| Deblinger<br>1996/1999 | Trauma-focused<br>CBT: Exposure<br>therapy/prolonged<br>exposure (PE) | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Childhood sexual abuse (Contact sexual abuse. 18% experienced 1 sexually abusive incident, 47% 2-10 episodes, 22% 11-50 episodes, and 13% >50 abusive incidents) | 100 | Age range (mean): 7-13(9.8) Gender (% female): 83 BME (% non-white): 28 Country: US Coexisting conditions: 29% major depression; 30% oppositional defiant disorder; 20% ADHD; 11% separation anxiety; 6% conduct disorder; 5% specific phobia; 1% OCD Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion criteria: contact childhood sexual abuse substantiated by an investigation conducted by the Division of Youth and Family Services (DYFS) or the prosecutor's office; children aged 7-13 years; consent to participate in the study completed by child and legal guardian; presence of a total of three posttraumatic stress symptoms including at least one symptom of avoidance or reexperiencing. Exclusion criteria: severe developmental delay; psychosis; ongoing, unsupervised contact with the alleged perpetrator; female caretaker not willing to participate; danger to themselves or others |

| Study ID                        | Intervention  | PTSD details   | Trauma type   | N   | Demographics  | Inclusion/Exclusion criteria  |
|---------------------------------|---|--|---|-----|---|---|
| de Roos 2017                    | Trauma-focused<br>CBT: Narrative<br>exposure therapy<br>(NET)           | Clinically important PTSD symptoms (scoring above a threshold on validated scale)                | Mixed - Physical abuse/assault (23%); Sexual abuse (26%); Accident/injury of a loved one (19%); Traumatic loss (18%); Disaster/other (13%)  | 103 | Age range (mean): 8-18 (13.1) Gender (% female): 57 BME (% non-white): NR Country: Netherlands Coexisting conditions: 54% had one or more co- morbid disorder (assessed with ADIS-C) Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion criteria: children aged 8-18 years; able to read/write and communicate in the Dutch language; have a primary DSM-IV diagnosis of PTSD or subthreshold PTSD, with the PTSD symptoms being tied to a single traumatic event that occurred at least one month prior to trial assessment. Exclusion criteria: the presence of symptoms other than PTSD in more urgent need of treatment (e.g., suicidal intent/acts, acute psychosis); ongoing exposure to a severe threat to the child's safety; starting psychotropic medication within three months of trial assessment; currently receiving another form of psychological treatment; an IQ estimated to be < 80 based on information contained in the medical history or referral letter. |
| Diehle<br>2015/Lindauer<br>2009 | Trauma-focused<br>CBT: Cohen TF-<br>CBT/Cognitive<br>processing therapy | Clinically<br>important PTSD<br>symptoms<br>(scoring above<br>a threshold on<br>validated scale) | Mixed - 63% Single-<br>event index trauma.<br>Single event traumas:<br>accidents (23 %),<br>sexual assault (17 %);<br>threat (with weapon)<br>(13 %); kidnapping (10<br>%); serious illness (7<br>%); or other (30 %).<br>Multiple-event traumas: | 48  | Age range (mean): 8- 18(12.9) Gender (% female): 62 BME (% non-white): NR Country: Israel Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with  | Inclusion criteria: aged 8-18 years; command of the Dutch language; exposure to at least one single traumatic event; the last traumatic event occurred at least 4 weeks prior to the first measurement; and partial (fulfilling two of the three symptom clusters or one symptom present in each of the three symptom clusters) or full PTSD as   |

| Study ID                    | Intervention   | PTSD details  | Trauma type   | N  | Demographics   | Inclusion/Exclusion criteria   |
|-----------------------------|--|---|---|----|--|--|
|                             |  |   | exposure to domestic violence (44 %) and sexual assault (39 %) and other (17 %)   |    | previous trauma): Mean<br>types of prior trauma 6.5<br>Single or multiple incident<br>index trauma: Single   | reported by the child (interviewed with the CAPS-CA) or the caretaker (interviewed with the ADIS-P PTSD module). Exclusion criteria: clinical signs of psychotic disorder, substance use disorder, pervasive developmental disorder (e.g., autism) or acute suicidality. After 12 months of slow recruitment, the inclusion criteria was adjusted in order to also include children who had experienced multiple-event trauma. |
| Ertl<br>2011/Neuner<br>2007 | Trauma-focused CBT: Narrative exposure therapy for traumatized children and adolescents (KidNET) | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Child soldiers - The duration of abduction ranged from several hours to 7.42 years, with a median of 2.47 months. Other than abduction, the most common traumatic event types reported by 81 or more of the 85 participants were exposure to a war zone, witnessing someone being killed, witnessing abduction, witnessing physical assault, and assaults with weapons. The likelihood of an event being indicated as the | 85 | Age range (mean): 12-25(18.4) Gender (% female): 55 BME (% non-white): NR Country: Uganda Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | Inclusion criteria: Former child soldiers aged 12-25 years, whose PTSD diagnoses were confirmed by clinical experts according to the Clinician- Administered PTSD Scale (CAPS). Exclusion criteria: psychotic symptoms.  |

| Study ID   | Intervention  | PTSD details  | Trauma type   | N  | Demographics  | Inclusion/Exclusion criteria  |
|--|---|---|---|----|---|---|
|  |   |   | worst if present was<br>highest for being forced<br>to kill (55%), followed<br>by witnessed killing<br>(31%) and seeing<br>someone being<br>mutilated or seeing<br>dead bodies (13%)                      |    |   |   |
| Foa<br>2013a/McLean<br>2015a/Capaldi<br>2016/Kaczkurkin<br>2016/Zandberg<br>2016 | Trauma-focused<br>CBT: Exposure<br>therapy/prolonged<br>exposure (PE)   | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Childhood sexual abuse  | 61 | Age range (mean): 13-18 (15.3) Gender (% female): 100 BME (% non-white): 82 Country: US Coexisting conditions: 57% had ≥1 comorbid psychiatric diagnoses Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | Inclusion criteria: adolescent girls who met criteria for a primary DSM-IV-TR diagnosis of chronic or subthreshold (required only two of three avoidance symptoms and a total score 14 on the Child Posttraumatic Stress Scale-Interview) PTSD related to sexual assault. Exclusion criteria: suicidal ideation with intent, current uncontrolled bipolar disorder, current psychosis, current conduct disorder, pervasive developmental disorder, and initiation of psychotropic medication within the previous 12 weeks |
| Ford 2012  | Trauma-focused<br>CBT: Cohen TF-<br>CBT/Cognitive<br>processing therapy | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Mixed - Trauma<br>exposure was<br>extensive, including<br>97% to a traumatic<br>accident, disaster, or<br>illness; 88% to physical<br>assault or abuse; 81%<br>to traumatic community<br>violence; 78% to | 59 | Age range (mean): 13-<br>17(14.7) Gender (% female): 100 BME (% non-white): 75 Country: US Coexisting conditions: 34% major depressive disorder, 26% oppositional defiant disorder, 23%   | Inclusion criteria: self-reported delinquency; full or partial PTSD (Clinician Administered PTSD Scale for Children-Adolescents [CAPS-CA] structured diagnostic interview). Exclusion criteria: substantial cognitive impairment (i.e., score <16 on Orientation, Attention, and Recall sections of   |

| Study ID                           | Intervention  | PTSD details  | Trauma type   | N   | Demographics  | Inclusion/Exclusion criteria  |
|------------------------------------|---|---|---|-----|---|---|
|                                    |   |   | traumatic family<br>violence; 44% to sexual<br>assault or abuse; 41%<br>to traumatic emotional<br>abuse; and 29% to<br>traumatic bullying |     | conduct disorder, and 13% attention deficit hyperactivity disorder Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple   | the Mini Mental State Exam; on one-to-one suicide watch (although suicidal ideation was not an exclusion, and most participants reported current or previous suicidal ideation); age younger than 13 or older than 18.  |
| Gilboa-<br>Schechtman<br>2004/2010 | Trauma-focused<br>CBT: Exposure<br>therapy/prolonged<br>exposure (PE) | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Mixed - Terrrorist attack (13%); motor vehicle accident (42%); non-sexual assault (0.5%); sexual assault (21%); Other (18%)               | 38  | Age range (mean): 12-18 (14.1) Gender (% female): 63 BME (% non-white): NR Country: Israel Coexisting conditions: 81% ≥ 1 comorbid disorder: 50% had one additional internalizing disorder, 13% had an additional externalizing disorder, and 16% had internalizing and externalizing disorders. Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion criteria: aged 12-18 years, a primary diagnosis of PTSD related to a single traumatic event, and fluency in Hebrew. Exclusion criteria: organic brain damage, mental retardation, an ongoing trauma-related threat, suicidal ideation posing imminent danger, current substance dependence, pending legal issues, initiation of treatment with psychotropic medication within the previous 6 weeks, and ongoing psychological treatment |
| Goldbeck<br>2016/Sachser<br>2016   | Trauma-focused CBT: Cohen TF-   | Clinically important PTSD symptoms  | Mixed - Interpersonal trauma (77%); accidental (23%). The   | 159 | Age range (mean): 7-17 (13)   | Inclusion criteria: aged 7–17 years; exposure to one or more traumatic event(s) after age 2 and dating  |

| Study ID | Intervention                     | PTSD details                                   | Trauma type   | N | Demographics   | Inclusion/Exclusion criteria  |
|----------|----------------------------------|--|---|---|--|---|
|          | CBT/Cognitive processing therapy | (scoring above a threshold on validated scale) | most frequently reported traumatic index events were experiences of sexual abuse, sexual assaults, physical violence, or witnessing domestic violence |   | Gender (% female): 72 BME (% non-white): NR Country: Germany Coexisting conditions: 34% >1 comorbid DSM-IV disorder: Depressive disorders (20%); Anxiety disorders (10%); ADHD (6%); Disruptive behaviour disorders (4%) Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Number of traumatic events: 6.35 (3.70) Single or multiple incident index trauma: Multiple | back at least 3 months; at least medium severity of PTSS as indicated by a total symptom score of ≥ 35 and at least one symptom per DSM-IV clusters B, C, and D assessed with the Clinician-Administered PTSD Scale for Children and Adolescents (CAPS-CA); PTSD was the main disorder based on clinical estimation, if comorbid disorders were present; availability of a non-offending adult caregiver for the treatment; willingness and ability of the patient and the caregiver to attend weekly treatment sessions; safe living circumstances to minimize the risk of re-traumatization during the study; sufficient cognitive ability to respond to cognitive interventions, as indicated by a raw score of ≥ 14 on the block design and vocabulary subtests of the Wechsler Intelligence Scale for Children (WISC IV); patients' and caregivers' sufficient command of the German language to participate in the treatment. Exclusion Criteria: acute suicidal behaviour or suicidal ideations requiring immediate hospitalization; severe head trauma indicated by a score <9 on the Glasgow Coma Scale as |

| Study ID    | Intervention                     | PTSD details  | Trauma type   | N  | Demographics   | Inclusion/Exclusion criteria  |
|-------------|----------------------------------|---|---|----|--|---|
|             |                                  |   |   |    |  | known from the patient's medical history, to avoid brain dysfunction or retrograde amnesia of the traumatic event due to head injury; a current or lifetime diagnosis of a pervasive developmental disorder or psychosis; psychopharmacological treatment started <6 weeks before recruitment or change of psychotropic medication during the course of the study; concurrent psychotherapy during the study; current severe mental disorder of the patient's main caregiver as evaluated by the responsible clinician, such as psychosis, severe episode of depression, or severe substance abuse; a sibling of the patient already participating in the study |
| Jaycox 2009 | Trauma-focused<br>CBT: CBT group | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Exposure to non-sexual violence (Experience of severe violence in the prior year) | 78 | Age range (mean): NR (11.5) Gender (% female): 51 BME (% non-white): 96 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion criteria: experience of severe violence in the prior year (responses on the Modified Life Experiences Survey [LES] indicating being the victim or witness of violence involving a knife of gun or having a summed score greater than 3, consistent with exposure to one or more violent events; current symptoms of PTSD as assessed on the Child PTSD Symptom Scale of 11 or   |

| Study ID            | Intervention  | PTSD details   | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria  |
|---------------------|---|--|--|-----|---|---|
|                     |   |  |  |     |   | greater, indicating moderate levels of symptom severity.  |
| Jensen<br>2014/2017 | Trauma-focused CBT: Cohen TF- CBT/Cognitive processing therapy        | Clinically important PTSD symptoms (scoring above a threshold on validated scale)                | Mixed - 59% violence or threats of violence outside the family context, 45.5% physical abuse within the family, 42.9% witnessing violence within the family, 27.6% witnessing violence outside the family, 27.6% sexual abuse outside the family, 20.5% severe accident, 16% extremely painful or frightening medical procedures, 10.9% robbery or assault, 7.7% sexual abuse within the family, 5.8% natural disaster, 5.1% kidnapping, and 30.8% other frightening or overwhelming experiences | 156 | Age range (mean): 10-18 (15.1) Gender (% female): 80 BME (% non-white): NR Country: Israel Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean 3.6 different types of traumas (SD=1.8, range=1–10) Single or multiple incident index trauma: Multiple | Inclusion criteria: children aged 10-18 years who had been referred to one of eight community mental health outpatient clinics in Norway; the youth had to have experienced at least one traumatizing event and suffered from significant posttraumatic stress reactions (score ≥ 15 on the CPSS and at least one symptom in each of the three PTSD symptom criteria [re-experiencing, avoidance, and hyperarousal]). Exclusion criteria: acute psychosis; suicidal behaviour; need for an interpreter. |
| King 2000           | Trauma-focused<br>CBT: Exposure<br>therapy/prolonged<br>exposure (PE) | Clinically<br>important PTSD<br>symptoms<br>(scoring above<br>a threshold on<br>validated scale) | Childhood sexual<br>abuse (In the majority<br>of cases, the offenders<br>were male adults<br>known to the child such<br>as the biological father,<br>stepfather, family  | 36  | Age range (mean): 5-17 (11.4) Gender (% female): 69 BME (% non-white): NR Country: Australia Coexisting conditions: For 69% who met DSM-IV  | Inclusion criteria: sexually abused children who had a history of contact sexual abuse such as sexual touching, oral-genital contact, or penile penetration of the vagina or anus, with or without physical force; if the perpetrator   |

| Study ID     | Intervention                  | PTSD details                       | Trauma type   | N  | Demographics   | Inclusion/Exclusion criteria  |
|--------------|-------------------------------|------------------------------------|---|----|--|---|
|              |                               |                                    | friend, neighbour, or teacher. Nearly all of the children had experienced multiple episodes of sexual abuse involving penetration offenses and other forms of sexual abuse) |    | criteria for full PTSD (N=25): 16% with full PTSD had no other Axis I diagnoses, 36% had one comorbid diagnosis, 40% had two comorbid diagnoses, and 8% had three comorbid diagnoses. The comorbid diagnoses included dysthymia (28%), oppositional defiant disorder (28%), separation anxiety disorder (24%), generalized anxiety disorder (20%), conduct disorder (12%), major depression (8%), attention-deficit/hyperactivity disorder (8%), and specific phobia (8%). Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean number of abusive episodes: 7.6 (SD=3.8; range 1-33) Single or multiple incident index trauma: Multiple | was a child, he/she had to be 5 years older than the victimized child; both child and his/her non- offending primary caregiver were English-speaking; independent validation of sexual abuse was required; the child's symptoms met diagnostic criteria for PTSD or fell short of the diagnostic criteria for PTSD by several symptoms, but the child still experienced severe stress reactions (had to exhibit a total of 3 posttraumatic stress symptoms including at least one of avoidance or re-experiencing phenomena). Exclusion criteria: ongoing, unsupervised contact with the alleged perpetrator; severe intellectual disability, psychosis, or suicidal behaviour; taking antidepressant or antianxiety medication; child or parents not willing to participate. |
| Langley 2015 | Trauma-focused CBT: CBT group | Clinically important PTSD symptoms | Mixed (Types of trauma commonly reported included: Witnessed/   | 74 | Age range (mean): 6-11 (7.7)   | Inclusion criteria: experience of one or more traumatic events; current symptoms of PTSD  |

| Study ID   | Intervention                     | PTSD details  | Trauma type   | N   | Demographics  | Inclusion/Exclusion criteria   |
|------------|----------------------------------|---|---|-----|---|--|
|            |                                  | (scoring above a threshold on validated scale)                                    | know of family member arrested (31%); Witnessed physical violence (26%); Victim of physical violence (25%); Witnessed or heard about neighbourhood or school violence (25%); Separated from parent(s) (e.g., deportation, deployment, hospitalization) (22%); Witnessed a serious accident (18%); Threatened by someone (violence) (18%); Someone close to child very sick or hurt badly (16%); Serious Illness/hospitalization of loved one (15%)) |     | Gender (% female): 50<br>BME (% non-white): 73<br>Country: US<br>Coexisting conditions: NR<br>Lifetime experience of<br>trauma (mean number of<br>prior traumas/% with<br>previous trauma): NR<br>Single or multiple incident<br>index trauma: Multiple | indicating moderate or higher levels of symptom severity (score ≥20 on the PTSD Reaction Index). Exclusion criteria: presence of a severe psychiatric disturbance (i.e., acute suicidal behaviour, current psychotic symptoms); sexual abuse as only and primary trauma.   |
| Layne 2008 | Trauma-focused<br>CBT: CBT group | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Witnessing war as a civilian (Approximately 73% of the students participating reported experiencing direct life threat arising from close proximity to exploding shells or rifle fire, 36% reported witnessing during the war violent death or  | 159 | Age range (mean): 13-19(16) Gender (% female): 64 BME (% non-white): NR Country: Bosnia Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR   | Three inclusion criteria: significant trauma exposure before, during, and/or after the war (e.g., serious physical injury, life threat, witnessing death or serious injury, traumatic bereavement); significant current distress, especially severe persisting symptoms of PTSD, depression, or traumatic grief; and significant functional impairment, including family or peer |

| Study ID                    | Intervention                                | PTSD details  | Trauma type  | N  | Demographics  | Inclusion/Exclusion criteria   |
|-----------------------------|---|---|--|----|---|--|
|                             |   |   | serious injury, 12% reported witnessing torture, and 46% reported the serious injury of a person to whom they were close, 14% reported the violent death during the war of a nuclear family member, and 73% reported the violent death of at least one person to whom they were close) |    | Single or multiple incident index trauma: Multiple  | relationships and school performance. With respect to exclusion criteria, students who did not meet the three inclusion criteria or who did but showed signs of psychosis, represented an imminent threat to themselves or others, were unable to attend group meetings, were judged not to be appropriate for group-based intervention due to highly disruptive behavioural or substance abuse problems, or reluctance to participate in a group setting were excluded from participation in the study  |
| Meiser-Stedman<br>2010/2017 | Trauma-focused<br>CBT: Cognitive<br>therapy | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Motor Vehicle<br>Collisions: Motor<br>vehicle collision (52%);<br>Assault (24%); Medical<br>emergency (3%);<br>House fire (3%); Other<br>(17%)   | 29 | Age range (mean): 8-17 (13.3) Gender (% female): 72 BME (% non-white): 14 Country: UK Coexisting conditions: 86% comorbid anxiety disorder; 55% comorbid affective disorder; 52% comorbid behavioural disorder Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): 38% had experienced previous trauma | Inclusion criteria: aged 8–17 years old; main presenting problem of PTSD (using an age-appropriate diagnostic algorithm [PTSD-AA]: presence of one re-experiencing symptom, one avoidance symptom, two hyperarousal symptoms and impaired functioning) relating to a single trauma in previous 2–6 months; fluency in English. Exclusion criteria: organic brain damage; unconscious >15 min during the trauma; intellectual disability or autistic spectrum disorder; ongoing threat; recently initiated (within 3 months) psychotropic medication; receiving another psychological |

| Study ID              | Intervention   | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria   |
|-----------------------|--|---|--|----|--|--|
|                       |  |   |  |    | Single or multiple incident index trauma: Single   | treatment; acute treatment required for suicide risk or other major mental health problem.   |
| Pityaratstian<br>2015 | Trauma-focused<br>CBT: Brief group<br>CBT  | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Natural disasters (such as severe floods, earthquakes or tsunamis): Tsunami in Thailand - 50% saw tsunami with own eyes; 36% lost family member; 64% lost friend; 25% lost home; 28% sustained injury  | 36 | Age range (mean): 10-15 (12.3) Gender (% female):72 BME (% non-white):NR Country: Thailand Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single  | Inclusion criterion: primary diagnosis of DSM-IV-TR PTSD. Exclusion criteria: mental retardation, pervasive developmental disorders, psychotic symptoms, or current involvement in psychopharmacological treatment |
| Ruf 2010              | Trauma-focused CBT: Narrative exposure therapy for traumatized children and adolescents (KidNET) | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Witnessing war as a civilian - Violent attacks against their parents or other family members at home (73%) were the most common trauma type reported. These assaults were mainly conducted by soldiers or other organized militant groups (58%). Other traumatic experiences included witnessing physical attacks against non-family members outside of the house (50%), accidents | 26 | Age range (mean): 7-16(11.4) Gender (% female): 46 BME (% non-white): NR Country: Germanyh Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Mean number of traumatic event types: 4.4 Single or multiple incident index trauma: Multiple | Children/young people aged 7-16, diagnosed with PTSD according to DSMIV, willing to participate and all parents giving consent. Exclusion: acute psychotic symptoms.   |

| Study ID   | Intervention                                | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria  |
|------------|---|---|--|----|--|---|
|            |   |   | (46%), violence against the child at home (35%, most of these were by militant forces, 27%), assaults against the child outside of the home (35%), living in a place of war (35%), seeing dead bodies (35%), painful or scary medical treatments (27%), hearing about the violent death of a beloved person (27%), earthquakes (19%), other natural disasters (12%), and sexual abuse (8%) |    |  |   |
| Smith 2007 | Trauma-focused<br>CBT: Cognitive<br>therapy | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Motor Vehicle<br>Collisions: Motor<br>vehicle accident (50%);<br>Assault (38%);<br>Witnessed violence<br>(13%)   | 24 | Age range (mean): NR (13.9) Gender (% female):50 BME (% non-white): 54 Country: UK Coexisting conditions: 79% had any comorbidity Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): 29% prior exposure to trauma Single or multiple incident index trauma: Single | Inclusion criteria: Children aged 8-18 years; presenting with PTSD symptoms relating to a single traumatic event; fluent in English; retained their PTSD diagnosis after 4 weeks of symptom monitoring. Exclusion criteria: organic brain damage; unconscious for >15 mins during trauma; significant learning difficulty; ongoing trauma-related threat in the environment; psychotropic medication (within 3 months); current other psychological treatment |

| Study ID                       | Intervention  | PTSD details  | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria  |
|--------------------------------|---|---|--|-----|---|---|
| Shein-Szydlo<br>2016           | Trauma-focused<br>CBT: Cohen TF-<br>CBT/Cognitive<br>processing therapy | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Mixed (Street Children in Mexico City - 56% were victims of sexual abuse, 47% of physical abuse, 18% of witnessing a violent event, and 17% of death of a family member) | 100 | Age range (mean): 12-18 (14.9) Gender (% female): 64 BME (% non-white): NR Country: Mexico Coexisting conditions: 14% anxiety disorder; 28% depression Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): 35% reported more than one type of traumatic event Single or multiple incident index trauma: Multiple | Inclusion criteria: children aged 12-18 years in one of eight facilities that provide shelter, food, basic education, and medical care for street children in Mexico City; with a PTS score of ≥24 at screening; had a diagnosis of PTSD (assessed with the Diagnostic Interview Schedule for Children). Exclusion criteria: severe psychopathology (psychosis, severe suicidal depression requiring different and immediate treatment); intellectual disability        |
| Stein<br>2003a/Kataoka<br>2011 | Trauma-focused<br>CBT: CBT group  | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Exposure to non-sexual violence (76% any violence involving a gun or knife. Number of violent events experienced: 2.8; Number of violent events witnessed: 5.95)         | 126 | Age range (mean): NR (11) Gender (% female): 56 BME (% non-white): NR Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple  | Inclusion criteria: Children aged 11-12 years with substantial exposure to violence (victom or witness) involving knife or gun or having a Life Events Scale score >6, consistent with exposure to≥3 violent events; clinically important PTSD symptoms on CPSS scale; PTSD symptoms related to exposure to violence that they were willing to discuss in a group; not disruptive/able to participate in intervention according to school-based mental health clinician |

ADHD-Attention deficit hyperactivity disorder; BME-Black and minority ethnic; CBT-Cognitive Behaviour Therapy; DSM-Diagnostic and Statistical Manual of Mental Disorders; ICD-International Classification of Disease; LED-Modified life experiences surveys; NET-Narrative exposure therapy; NR-Not recorded; PTSD-Post-traumatic stress disorder; PTSS-Post-traumatic stress syndrome.

### Psychological: Non-trauma-focused CBT

| Study ID      | Intervention  | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria  |
|---------------|---|---|--|----|--|---|
| Najavits 2006 | Non-trauma-<br>focused<br>CBT:<br>Seeking<br>Safety | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Mixed - The most common trauma category was sexual abuse (88%), followed by general disaster/accident (82%), physical abuse (73%), and crime (39%) | 33 | Age range (mean): NR (16.1) Gender (% female): 100 BME (% non-white): 21 Country: US Coexisting conditions: All met current DSM-IV criteria for both PTSD and SUD, with 94% having substance dependence. Current substance dependence diagnoses per DSM-IV criteria at intake were: cannabis (79%), alcohol (67%), hallucinogens | Inclusion criteria: outpatient adolescent girls; met current DSM-IV criteria for both PTSD and SUD; active substance use within the past 60 days. Exclusion criteria: a history of bipolar I disorder, psychotic disorder, were mandated to treatment, or had characteristics that would interfere with treatment completion (mental retardation, homelessness, impending incarceration, or a life-threatening illness) |

| Study ID Interven | ention PTSD details | Trauma type | N | Demographics   | Inclusion/Exclusion criteria |
|-------------------|---------------------|-------------|---|--|------------------------------|
|                   |                     |             |   | (21%), amphetamines (15%), cocaine (9%), opioids (9%), inhalants (9%), barbiturates (6%), polysubstance (6%), and PCP 1 (3%). Participants could have more than one diagnosis Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple |                              |

DSM-Diagnostic and Statistical Manual of Mental Disorders; ICD-International Classification of Disease; NR-Not recorded; PTSD-Post-traumatic stress disorder; SUD-Substance use disorder

**Psychological: Supportive counselling** 

| Study ID                    | Intervention   | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria   |
|-----------------------------|--|---|--|----|--|--|
| Chen 2014                   | Trauma-<br>focused<br>CBT: CBT<br>group  | Clinically important<br>PTSD symptoms<br>(scoring above a<br>threshold on<br>validated scale)     | Natural disasters (such as severe floods, earthquakes or tsunamis): Adolescents who had lost at least 1 parent in the Sichuan, China, Earthquake   | 40 | Age range (mean): NR (14.5) Gender (% female): 68 BME (% non-white): NR Country: Israel Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single | Inclusion criteria: adolescents from two secondary schools, who had lost at least one parent in the earthquake, and scored≥18 on the CRIES-13  |
| Ertl<br>2011/Neuner<br>2007 | Trauma-<br>focused<br>CBT:<br>Narrative<br>exposure<br>therapy for<br>traumatized<br>children and<br>adolescents<br>(KidNET) | PTSD diagnosis<br>according to<br>ICD/DSM criteria<br>(including self-<br>report of<br>diagnosis) | Child soldiers - The duration of abduction ranged from several hours to 7.42 years, with a median of 2.47 months. Other than abduction, the most common traumatic event types reported by 81 or more of the 85 participants were exposure to a war zone, | 85 | Age range<br>(mean): 12-<br>25(18.4)<br>Gender (%<br>female): 55<br>BME (% non-<br>white): NR<br>Country:<br>Uganda  | Inclusion criteria: Former child soldiers aged 12-25 years, whose PTSD diagnoses were confirmed by clinical experts according to the Clinician-Administered PTSD Scale (CAPS). Exclusion criteria: psychotic symptoms. |

| Study ID | Intervention | PTSD details | Trauma type  | N | Demographics   | Inclusion/Exclusion criteria |
|----------|--------------|--------------|--|---|--|------------------------------|
|          |              |              | witnessing someone being killed, witnessing abduction, witnessing physical assault, and assaults with weapons. The likelihood of an event being indicated as the worst if present was highest for being forced to kill (55%), followed by witnessed killing (31%) and seeing someone being mutilated or seeing dead bodies (13%) |   | Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple |                              |

BME-Black and minority ethnic; CAPS- Clinician administered PTSD scale; CBT-Cognitive Behaviour Therapy; DSM-Diagnostic and Statistical Manual of Mental Disorders; ICD-International Classification of Disease; NR-Not recorded; PTSD-Post-traumatic stress disorder;

#### Psychological: Eye movement desensitisation and reprocessing (EMDR)

| Study ID           | Intervention  | PTSD details  | Trauma type  | N  | Demographics  | Inclusion/Exclusion criteria  |
|--------------------|---------------|---|--|----|---|---|
| Ahmad<br>2007/2008 | EMDR:<br>EMDR | PTSD diagnosis<br>according to<br>ICD/DSM criteria<br>(including self-<br>report of<br>diagnosis) | Mixed - Maltreatment (36.4%), sexual abuse (21.2%), road accident (15.2%), witnessing unnatural death (12.1%) and other types of trauma (6.1%) | 33 | Age range (mean):<br>6-16 (9.9)<br>Gender (% female):<br>61<br>BME (% non-white):<br>NR<br>Country: Sweden<br>Coexisting<br>conditions: 79%<br>fulfilled DSM-IV | Inclusion criteria: a PTSD diagnosis, child aged at least 6 years, no manifest learning disabilities, experienced at least one traumatic experience, and grown up in at least one socially exposed condition (defined as the child having grown up with a family member with criminality, substance abuse, chronic illness, handicap, or having the caregiver physically or mentally unavailable for the child). Exclusion criteria: if children needed other types of treatment (such as medication, |

| Study ID     | Intervention  | PTSD details  | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria  |
|--------------|---|---|--|-----|---|---|
|              |   |   |  |     | criteria for at least one additional diagnosis: Depression (46%); ADHD (30%); ODD (21%); separation anxiety (18%); conduct disorder (12%), overanxious disorder and autism spectrum (3%) Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | cognitive therapy or play therapy), or received social welfare service during the study   |
| de Roos 2017 | Trauma-<br>focused<br>CBT:<br>Narrative<br>exposure<br>therapy<br>(NET) | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Mixed - Physical<br>abuse/assault (23%);<br>Sexual abuse (26%);<br>Accident/injury of a loved<br>one (19%); Traumatic<br>loss (18%);<br>Disaster/other (13%) | 103 | Age range (mean): 8-18 (13.1) Gender (% female): 57 BME (% non-white): NR Country: Netherlands Coexisting conditions: 54% had one or more co- morbid disorder   | Inclusion criteria: children aged 8-18 years; able to read/write and communicate in the Dutch language; have a primary DSM-IV diagnosis of PTSD or subthreshold PTSD, with the PTSD symptoms being tied to a single traumatic event that occurred at least one month prior to trial assessment. Exclusion criteria: the presence of symptoms other than PTSD in more urgent need of treatment (e.g., suicidal intent/acts, acute psychosis); ongoing exposure to a severe threat to the child's safety; starting psychotropic medication within three months of trial assessment; currently receiving another form of |

| Study ID         | Intervention  | PTSD details  | Trauma type                       | N  | Demographics  | Inclusion/Exclusion criteria   |
|------------------|---------------|---|-----------------------------------|----|---|--|
|                  |               |   |                                   |    | (assessed with ADIS-C) Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single   | psychological treatment; an IQ estimated to be < 80 based on information contained in the medical history or referral letter.  |
| Soberman<br>2002 | EMDR:<br>EMDR | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Unclear (Not reported in details) | 29 | Age range (mean): 10-16 (NR) Gender (% female): 0 BME (% non-white): NR Country: US Coexisting conditions: Other primary diagnoses included: Conduct Disorder (59%); Attention Deficit Hyperactive Disorder (17%), Learning Disability (14%), Substance Abuse (13%), and Oppositional/Defiant Disorder (3%) Lifetime experience of trauma (mean | Participants were included if they were: (1) boys with conduct problems in residential or day treatment. Participants were excluded if they: (1) had psychosis; (2) had suicidal or homicidal ideation; (3) had epilepsy; (4) had medical instability; (5) had low motivation to participate (3 or lower on a 0-10 scale); (6) were unable to identify a sufficiently traumatic memory (rating of 4 or higher on the 0-10 Subjective Units of Distress Scale (SUDS)) |

| Study ID | Intervention | PTSD details | Trauma type | N | Demographics  | Inclusion/Exclusion criteria |
|----------|--------------|--------------|-------------|---|---|------------------------------|
|          |              |              |             |   | number of prior<br>traumas/% with<br>previous trauma):<br>NR<br>Single or multiple<br>incident index<br>trauma: Unclear |                              |

BME-Black and minority ethnic; DSM-Diagnostic and statistical manual of mental disorders; EMDR-Eye movement desensitisation and reprocessing; NR-Not recorded; PTSD-Post-traumatic stress disorder; NET-Narrative exposure therapy; SUDS-Subjective units of distress scale.

#### Psychological: Parent training/family interventions

| Study ID               | Intervention  | PTSD details  | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria  |
|------------------------|---|---|--|-----|---|---|
| Deblinger<br>1996/1999 | Trauma-<br>focused<br>CBT:<br>Exposure<br>therapy/prol<br>onged<br>exposure<br>(PE) | Clinically important<br>PTSD symptoms<br>(scoring above a<br>threshold on<br>validated scale) | Childhood sexual abuse<br>(Contact sexual abuse.<br>18% experienced 1<br>sexually abusive incident,<br>47% 2-10 episodes, 22%<br>11-50 episodes, and 13%<br>>50 abusive incidents) | 100 | Age range (mean): 7- 13(9.8) Gender (% female): 83 BME (% non-white): 28 Country: US Coexisting conditions: 29% major depression; 30% oppositional defiant disorder; 20% ADHD; 11% separation anxiety; 6% | Inclusion criteria: contact childhood sexual abuse substantiated by an investigation conducted by the Division of Youth and Family Services (DYFS) or the prosecutor's office; children aged 7-13 years; consent to participate in the study completed by child and legal guardian; presence of a total of three posttraumatic stress symptoms including at least one symptom of avoidance or re-experiencing. Exclusion criteria: severe developmental delay; psychosis; ongoing, unsupervised contact with the alleged perpetrator; female caretaker not willing to participate; danger to themselves or others |

| Study ID   | Intervention                                     | PTSD details  | Trauma type  | N   | Demographics  | Inclusion/Exclusion criteria   |
|------------|--|---|--|-----|---|--|
|            |  |   |  |     | conduct disorder; 5% specific phobia; 1% OCD Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Single                   |  |
| Kazak 2004 | Family<br>therapy:<br>Family<br>therapy<br>group | Clinically important<br>PTSD symptoms<br>(scoring above a<br>threshold on<br>validated scale) | Diagnosis of life-<br>threatening condition -<br>Diagnoses included<br>leukaemia (25%), solid<br>tumours (22%), lymphoma<br>(21%), bone tumours<br>(8%), and other (24%) | 150 | Age range (mean): 10-19(14.6) Gender (% female): 52 BME (% non-white): 12 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR | Participants were included if they were: (1) childhood cancer survivors aged 11-19 years; (2) had completed treatment 1–10 years previously; (3) on the oncology tumour registry. Participants were excluded if they: (1) experienced a relapse; (2) had mental retardation; (3) were not fluent in English; (4) resided more than 150 miles from the hospital |

| Study ID | Intervention | PTSD details | Trauma type | N | Demographics  | Inclusion/Exclusion criteria |
|----------|--------------|--------------|-------------|---|---|------------------------------|
|          |              |              |             |   | Single or<br>multiple incident<br>index trauma:<br>Single |                              |

ADHD-Attention deficit hyperactivity disorder; BME-Black and minority ethnic; CBT-Cognitive Behaviour Therapy; DYFS-Division of Youth and Family Services; NR-Not recorded; OCD-Obsessive Compulsive Disorder; PE-Prolonged exposure; PTSD-Post-traumatic stress disorder.

#### **Psychological: Combined somatic and cognitive therapies**

| Study ID           | Intervention   | PTSD details   | Trauma type                       | N  | Demographics  | Inclusion/Exclusion criteria  |
|--------------------|--|--|-----------------------------------|----|---|---|
| Al-Hadethe<br>2015 | Trauma-focused<br>CBT: Narrative<br>exposure therapy<br>for traumatized<br>children and<br>adolescents<br>(KidNET) | PTSD diagnosis according to ICD/DSM criteria (including self- report of diagnosis) | Unclear (Not reported in details) | 60 | Age range (mean): 16-19 (NR) Gender (% female): 0 BME (% non- white): Unclear Country: Iraq Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Unclear | Inclusion criteria: male secondary school students aged 16-19 years old, who met DSM-IV criteria for PTSD as measured by the Scale of Posttraumatic Stress Symptoms (SPTSS) |

BME-Black and minority ethnic; CBT-Cognitive Behaviour Therapy; DSM-Diagnostic and Statistical Manual of Mental Disorders; ICD-International Classification of Disease; NR-Not reported; PTSD-Post-traumatic stress disorder.

#### **Psychological: Play therapy**

| Study ID                                   | Intervention   | PTSD details  | Trauma type   | N    | Demographics   | Inclusion/Exclusion criteria   |
|--|--|---|---|------|--|--|
| Deeba 2015                                 | Play therapy:<br>Play therapy                                | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Mixed - Most of the children (90%) had lost one or both parents following natural disasters or accidents or due to domestic violence and witnessed direct or indirect violence against a parent (mostly towards the mother) | 13 1 | Age range (mean): 5-9 (7.2) Gender (% female): 37 BME (% non-white): NR Country: Bangladesh Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | Inclusion criteria: aged 5-9 years; lived in one of two shelter homes (children were accepted to both institutions following loss or abandonment by their parents and an absence of relatives able or willing to care for them); witnessed or experienced at least one severe DSM-IV traumatic event; provided assent to participate. Exclusion criteria: serious health conditions; psychotic features; severe ADHD; any developmental disorders; inability to comprehend simple instructions |
| Lieberman<br>2005/2006/Ghosh<br>Ippen 2011 | Play therapy:<br>Child-Parent<br>Psychotherapy<br>using play | Clinically important PTSD symptoms (scoring above a                               | Domestic violence (not reported in details)   | 75   | Age range (mean): 3-5 (4.1)  | Child–mother dyads were recruited if the child was 3 to 5 years old, had been exposed to marital violence as confirmed by mother's report on the Conflict Tactics Scale 2 (Straus 1996),   |

| Study ID | Intervention | PTSD details                  | Trauma type | N | Demographics   | Inclusion/Exclusion criteria  |
|----------|--------------|-------------------------------|-------------|---|--|---|
|          |              | threshold on validated scale) |             |   | Gender (% female): 52 BME (% non-white): 91 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): Multiple stressors, including exposure to community violence (46.7%), physical abuse (18.7%), sexual abuse (14.7%), or both (4%). During the study, 33.3% of the mothers reported new traumas that affected the dyad and 17.3% of the mothers | and the perpetrator was not living in the home. Exclusionary criteria for the mothers were documented abuse of the target child, current substance abuse and homelessness, mental retardation, and psychosis. Children with mental retardation or autistic spectrum disorder were also excluded |

| Study ID          | Intervention                  | PTSD details  | Trauma type   | N  | Demographics   | Inclusion/Exclusion criteria   |
|-------------------|-------------------------------|---|---|----|--|--|
|                   |                               |   |   |    | reported either returning to their violent partners or entering a new violent relationship Single or multiple incident index trauma: Multiple  |  |
| Schottelkorb 2012 | Play therapy:<br>Play therapy | Clinically important PTSD symptoms (scoring above a threshold on validated scale) | Witnessing war as a civilian (Childhood Refugee Trauma) | 31 | Age range (mean): 6-13 (9.2) Gender (% female): 45 BME (% non-white): 67 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index | Inclusion criteria: refugee children at one of three elementary schools; met criteria for full or partial PTSD assessed with the UCLA PTSD Index for DSM–IV or a score in the clinical range on the Parent Report of Posttraumatic Symptoms. Exclusion criteria: participating in counselling outside of the study |

| Study ID | Intervention | PTSD details | Trauma type | N | Demographics        | Inclusion/Exclusion criteria |
|----------|--------------|--------------|-------------|---|---------------------|------------------------------|
|          |              |              |             |   | trauma:<br>Multiple |                              |

ADHD-Attentions deficit hyperactivity disorder; DSM-Diagnostic and Statistical Manual of Mental Disease; NR-Not recorded; PTSD-Post-traumatic stress disorder.

#### **Psychosocial: Art therapy**

| Study ID               | Intervention  | PTSD details  | Trauma type  | N  | Demographics  | Inclusion/Exclusion criteria   |
|------------------------|---|---|--|----|---|--|
| Lyshak-Stelzer<br>2007 | Art therapy:<br>Trauma-<br>focused<br>expressive<br>art therapy | Clinically important<br>PTSD symptoms<br>(scoring above a<br>threshold on<br>validated scale) | Mixed (Frequently reported trauma types included: Physically abused or threatened with physical abuse at home (62%); Witnessing physical abuse at home (50%); Being in a bad accident (50%); Witnessing shooting, beating, or threats in neighbourhood (47%); Sexual abuse (46%); Beaten, shot at, or threatened in neighbourhood (45%); Serious medical problem (40%); Being in a disaster (weather, fire, etc.) (19%)) | 77 | Age range (mean): 13-17 (15.1) Gender (% female): 45 BME (% non-white): 82 Country: US Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | Inclusion criteria: participants aged 13-18 years; were able to sustain a school programme for 2 weeks running; were expected to stay at the hospital for at least 16 weeks from the date of parent or guardian consent. Exclusion criteria (at the Long Island facility only): Participants who were court-mandated |

NR-Not reported; PTSD-Post-traumatic stress disorder.

#### **Psychosocial: Meditation**

| Study ID            | Intervention                             | PTSD details  | Trauma type  | N  | Demographics   | Inclusion/Exclusion criteria  |
|---------------------|--|---|--|----|--|---|
| Gordon<br>2006/2008 | Meditation:<br>Mind-body<br>skills group | PTSD diagnosis according to ICD/DSM criteria (including self-report of diagnosis) | Witnessing war as a civilian (Kosovar adolescents) | 82 | Age range (mean): 14- 18(16.3) Gender (% female): 76 BME (% non- white): NR Country: Kosovo Coexisting conditions: NR Lifetime experience of trauma (mean number of prior traumas/% with previous trauma): NR Single or multiple incident index trauma: Multiple | Students at the high school under investigation who met criteria for PTSD (defined by meeting a threshold on the Albanian translation of the HTQ) |

DSM-Diagnostic and Statistical Manual of Mental Disorders; ICD-International Classification of Disease; NR-Not reported; PTSD-Post-traumatic stress disorder.

#### Appendix E – Forest plots

Forest plots for "For children and young people with clinically important posttraumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

Psychological: Trauma-focused CBT

Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD

Figure 2: Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at 1-month follow-up (CPTS-RI change score)

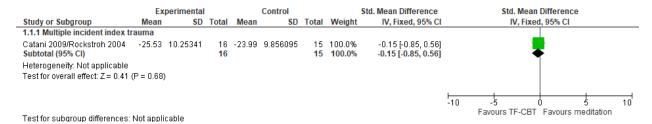


Figure 3: Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at 6-month follow-up (CPTS-RI change score)

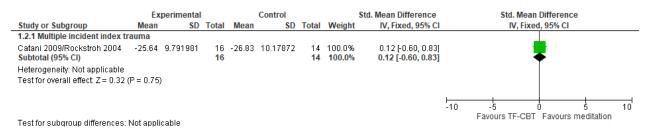


Figure 4: Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD: Diagnosis at 1-month follow-up (number of people who met criteria for a diagnosis of PTSD)

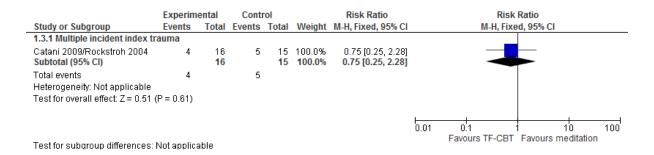
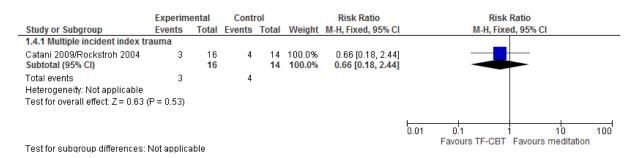


Figure 5: Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD: Diagnosis at 6-month follow-up (number of people who met criteria for a diagnosis of PTSD)



Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 6: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

## symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

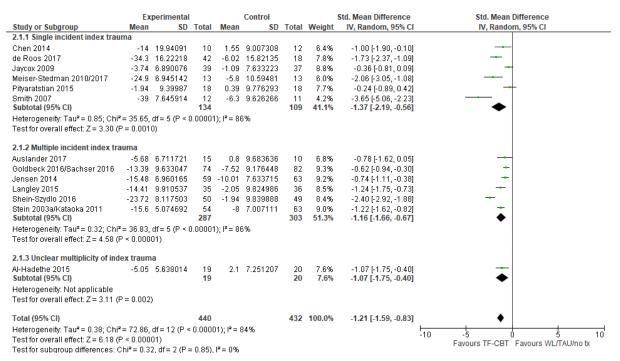


Figure 7: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 1-3 month follow-up (IES/SPTSS/CRIES/UCLA PTSD-RI/CPTS-RI change score)

|                                   | E            | kperimental               |           |           | Control           |       |        | Std. Mean Difference | Std. Mean Difference                                |
|-----------------------------------|--------------|---------------------------|-----------|-----------|-------------------|-------|--------|----------------------|---|
| Study or Subgroup                 | Mean         | <b>S</b> D                | Total     | Mean      | SD                | Total | Weight | IV, Random, 95% CI   | IV, Random, 95% CI                                  |
| 2.2.1 Single incident             | index tr     | auma                      |           |           |                   |       |        |                      |   |
| Berger 2009                       | -8.73        | 5.820653                  | 84        | -1.52     | 5.204805          | 82    | 33.1%  | -1.30 [-1.63, -0.96] | <b>+</b>  |
| Chen 2014                         | -22.8        | 8.747571                  | 10        | -2.2      | 9.071323          | 12    | 10.2%  | -2.22 [-3.33, -1.11] | <del></del>   |
| Pityaratstian 2015                | -5.67        | 8.496364                  | 18        | 0.78      | 10.15063          | 18    | 19.6%  | -0.67 [-1.35, 0.00]  | _=  |
| Subtotal (95% CI)                 |              |                           | 112       |           |                   | 112   | 62.9%  | -1.28 [-1.93, -0.63] | <b>◆</b>  |
| Heterogeneity: Tau² =             |              |                           |           | = 0.05)   | ; I² = 66%        |       |        |                      |   |
| Test for overall effect           | : Z = 3.88   | P = 0.0001                | 1)        |           |                   |       |        |                      |   |
| 2.2.2 Multiple incide             | nt index t   | trauma                    |           |           |                   |       |        |                      |   |
| Ahrens 2002                       | -12.11       | 8.049745                  | 19        | 0.08      | 5.759306          | 19    | 17.3%  | -1.71 [-2.46, -0.95] |   |
| Subtotal (95% CI)                 |              |                           | 19        |           |                   | 19    | 17.3%  | -1.71 [-2.46, -0.95] | <b>◆</b>  |
| Heterogeneity: Not ap             | pplicable    | 1                         |           |           |                   |       |        |                      |   |
| Test for overall effect           | Z = 4.43     | (P < 0.0000               | 01)       |           |                   |       |        |                      |   |
| 2.2.3 Unclear multip              | licity of in | ndex traum                | a         |           |                   |       |        |                      |   |
| Al-Hadethe 2015                   | -4           | 7.715646                  | 19        | 3.5       | 7.410408          | 20    | 19.8%  | -0.97 [-1.64, -0.30] |   |
| Subtotal (95% CI)                 |              |                           | 19        |           |                   | 20    | 19.8%  | -0.97 [-1.64, -0.30] | <b>◆</b>  |
| Heterogeneity: Not ap             | pplicable    | !                         |           |           |                   |       |        |                      |   |
| Test for overall effect           | Z = 2.85     | (P = 0.004)               |           |           |                   |       |        |                      |   |
| Total (95% CI)                    |              |                           | 150       |           |                   | 151   | 100.0% | -1.28 [-1.68, -0.87] | <b>•</b>  |
| Heterogeneity: Tau <sup>2</sup> = | = 0.10; CI   | $hi^2 = 7.90, dt$         | f= 4 (P   | = 0.10);  | I²= 49%           |       |        |                      | 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1               |
| Test for overall effect           | : Z = 6.16   | i (P < 0.0000             | 01)       |           |                   |       |        |                      | -10 -5 0 5 1<br>Favours TF-CBT Favours WL/TAU/no tx |
| Test for subgroup dif             | ferences     | : Chi <sup>2</sup> = 2.04 | l. df = 2 | (P = 0.3) | 36), $I^2 = 1.99$ | X6    |        |                      | TAVOUIS IT-ODT FAVOUIS WE/TAO/IIO K                 |

Figure 8: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 6-month follow-up (SPTSS change score)

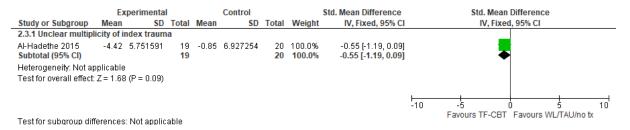


Figure 9: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 12-month follow-up (SPTSS change score)

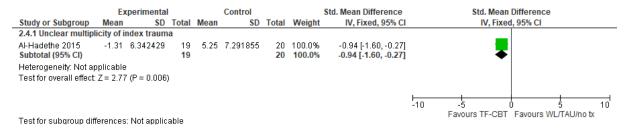


Figure 10: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score)

|  | Fx        | perimental     |           |                        | Control  |       |        | Std. Mean Difference |     | Std. Mean Difference               |    |
|--|-----------|----------------|-----------|------------------------|----------|-------|--------|----------------------|-----|------------------------------------|----|
| Study or Subgroup  | Mean      |                |           | Mean                   |          | Total | Weight | IV, Random, 95% CI   |     | IV, Random, 95% CI                 |    |
| 2.5.1 Single incident index trau                         | ma        |                |           |                        |          |       |        | ,                    |     |                                    |    |
| Meiser-Stedman 2010/2017                                 | -9.3      | 2.433105       | 13        | -1.5                   | 2.862691 | 13    | 11.1%  | -2.84 [-3.98, -1.71] |     | <b></b>                            |    |
| Smith 2007   | -48.9     | 12.01499       | 12        | -14.4                  | 12.1359  | 12    | 10.8%  | -2.76 [-3.93, -1.59] |     | <del></del>                        |    |
| Subtotal (95% CI)  |           |                | 25        |                        |          | 25    | 21.8%  | -2.80 [-3.62, -1.99] |     | •                                  |    |
| Heterogeneity: Tau² = 0.00; Chi²                         | = 0.01, d | lf = 1 (P = 0. | 92); l² = | = 0%                   |          |       |        |                      |     |                                    |    |
| Test for overall effect: $Z = 6.73$ (F                   |           |                |           |                        |          |       |        |                      |     |                                    |    |
| 2.5.2 Multiple incident index tra                        | uma       |                |           |                        |          |       |        |                      |     |                                    |    |
| Deblinger 1996/1999                                      | -5.48     | 2.118301       | 21        | -3.29                  | 2.339519 | 14    | 15.2%  | -0.97 [-1.69, -0.25] |     |                                    |    |
| Goldbeck 2016/Sachser 2016                               | -26.35    | 17.33886       | 76        | -14.1                  | 16.91013 | 83    | 19.0%  | -0.71 [-1.03, -0.39] |     | +                                  |    |
| Jensen 2014  | -29.64    | 16.75992       | 55        | -18.6                  | 17.62647 | 61    | 18.6%  | -0.64 [-1.01, -0.26] |     | +                                  |    |
| King 2000  | -5.75     | 3.007358       | 12        | -1.47                  | 1.681279 | 12    | 12.7%  | -1.70 [-2.65, -0.74] |     | <del></del>                        |    |
| Ruf 2010   | -26.1     | 9.750897       | 12        | -4.5                   | 12.33937 | 13    | 12.7%  | -1.87 [-2.84, -0.90] |     | <del></del>                        |    |
| Subtotal (95% CI)  |           |                | 176       |                        |          | 183   | 78.2%  | -0.98 [-1.37, -0.59] |     | <b>◆</b>                           |    |
| Heterogeneity: Tau <sup>2</sup> = 0.10; Chi <sup>2</sup> | = 9.27, d | f = 4 (P = 0.  | 05); l² = | = 57%                  |          |       |        |                      |     |                                    |    |
| Test for overall effect: $Z = 4.94$ (F                   | o.000     | 01)            |           |                        |          |       |        |                      |     |                                    |    |
| Total (95% CI)   |           |                | 201       |                        |          | 208   | 100.0% | -1.47 [-2.03, -0.90] |     | •                                  |    |
| Heterogeneity: Tau <sup>2</sup> = 0.41; Chi <sup>2</sup> | = 30.42.  | df = 6 (P < 1  | 0.00013   | ): I <sup>z</sup> = 80 | 1%       |       |        |                      | -   |                                    |    |
| Test for overall effect: $Z = 5.11$ (F                   |           |                | ,         |                        |          |       |        |                      | -10 | -5 0 5                             | 10 |
| Test for subgroup differences: C                         |           |                | < 0.00    | i01). i² =             | 93.6%    |       |        |                      |     | Favours TF-CBT Favours WL/TAU/no b | Į. |

Figure 11: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

## symptomatology clinician-rated at follow-up (CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score); multiple incident index trauma

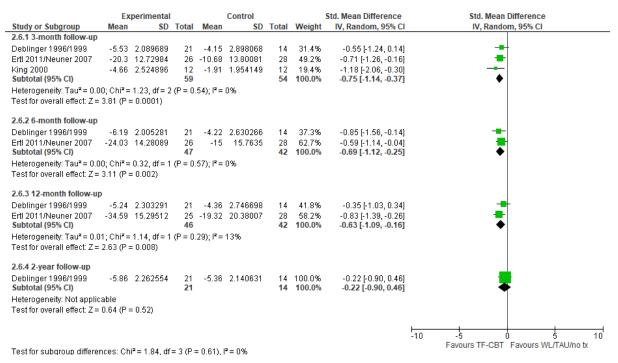


Figure 12: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at endpoint (number of people no longer meeting diagnostic criteria for PTSD)

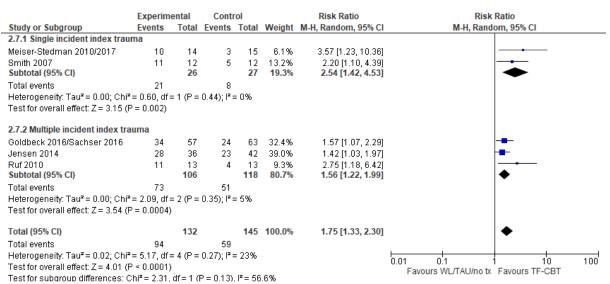


Figure 13: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at

### 1-3 month follow-up (number of people no longer above threshold on a scale for PTSD or meeting diagnostic criteria for PTSD)

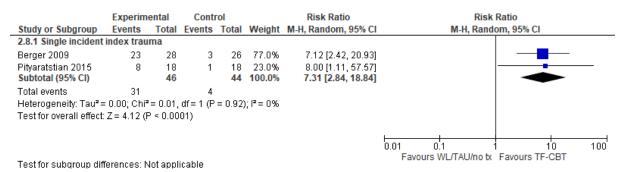


Figure 14: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at 12-month follow-up (number of people no longer meeting diagnostic criteria for PTSD)

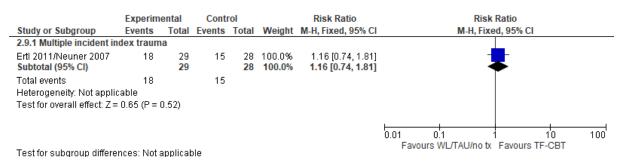


Figure 15: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Response at endpoint (number of people showing clinically significant improvement, based on reliable change indices [RCI]/rated as 'much/very much improved' on CGI)

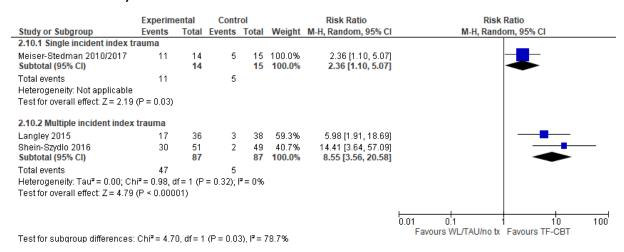


Figure 16: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms at endpoint (HADS-A/SCARED/RCMAS/SCAS/BAI change score)

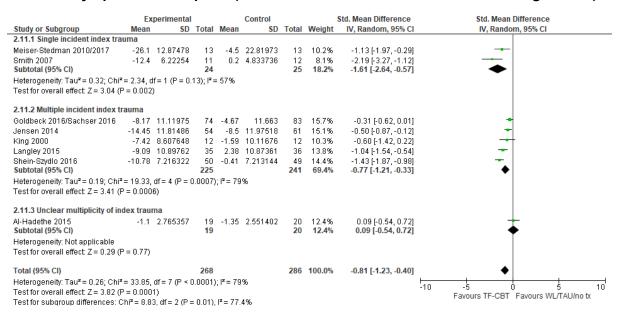


Figure 17: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms at 3-month follow-up (HADS-A/RCMAS change score)

|   | Ex           | perimental              | ı               |         | Control        |                 |                       | Std. Mean Difference                               | Std. Mean Difference                                |
|---|--------------|-------------------------|-----------------|---------|----------------|-----------------|-----------------------|--|---|
| Study or Subgroup                                 | Mean         | SD                      | Total           | Mean    | SD             | Total           | Weight                | IV, Random, 95% CI                                 | IV, Random, 95% CI                                  |
| 2.12.1 Multiple incide                            | ent index    | trauma                  |                 |         |                |                 |                       |  |   |
| King 2000<br>Subtotal (95% CI)                    | -9.92        | 9.885626                | 12<br><b>12</b> | -1.59   | 9.807839       | 12<br><b>12</b> | 44.6%<br><b>44.6%</b> | -0.82 [-1.66, 0.02]<br>- <b>0.82 [-1.66, 0.02]</b> | •   |
| Heterogeneity: Not ap<br>Test for overall effect: |              | (P = 0.06)              |                 |         |                |                 |                       |  |   |
| 2.12.2 Unclear multip                             | plicity of i | index traur             | ma              |         |                |                 |                       |  |   |
| Al-Hadethe 2015<br>Subtotal (95% CI)              | -1.42        | 2.842956                | 19<br><b>19</b> | -1.55   | 2.661128       | 20<br><b>20</b> | 55.4%<br><b>55.4%</b> | 0.05 [-0.58, 0.67]<br><b>0.05 [-0.58, 0.67]</b>    | <b>‡</b>  |
| Heterogeneity: Not ap<br>Test for overall effect  |              | (P = 0.89)              |                 |         |                |                 |                       |  |   |
| Total (95% CI)                                    |              |                         | 31              |         |                | 32              | 100.0%                | -0.34 [-1.18, 0.50]                                | •   |
| Heterogeneity: Tau <sup>2</sup> =                 | = 0.23; Ch   | i²= 2.60, d             | f=1 (P          | = 0.11) | ); I² = 62%    |                 |                       |  | 10 + 10   |
| Test for overall effect                           | Z = 0.79     | (P = 0.43)              |                 |         |                |                 |                       |  | 10 -5 0 5 10<br>Favours TF-CBT Favours WL/TAU/no tx |
| Test for subgroup dif                             | ferences:    | Chi <sup>2</sup> = 2.60 | 0, df = 1       | (P = 0. | .11), I² = 61. | 6%              |                       |  | FAVOUIS IF-CD1 FAVOUIS WE/TAO/IIO K                 |

Figure 18: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms at 6-month follow-up (HADS-A change score)

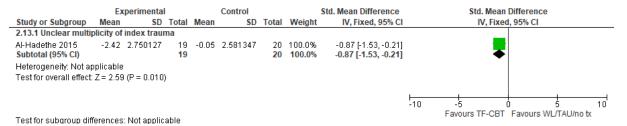


Figure 19: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms at 12-month follow-up (HADS-A change score)

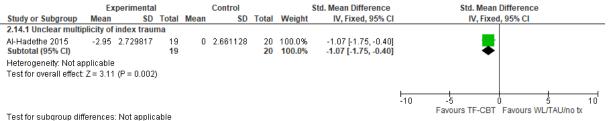


Figure 20: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at endpoint (HADS-D/CES-D/CDI/MFQ/DSRS/BDI change score)

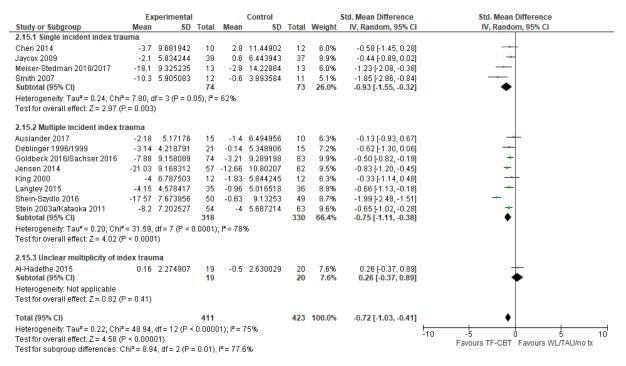


Figure 21: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 1-3 month follow-up (BDI/HADS-D/CES-D/CDI/MINI:Depression /DSRS change score)

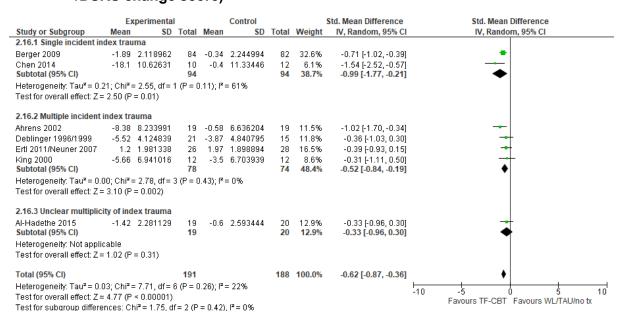


Figure 22: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 6-month follow-up (HADS-D/CDI/MINI:Depression change score)

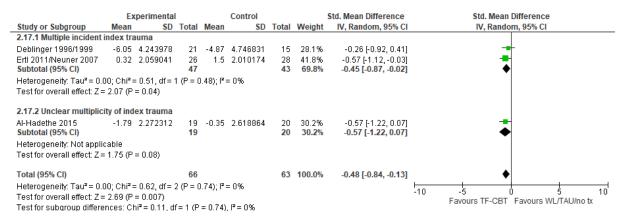


Figure 23: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 12-month follow-up (HADS-D/CDI/MINI:Depression change score)

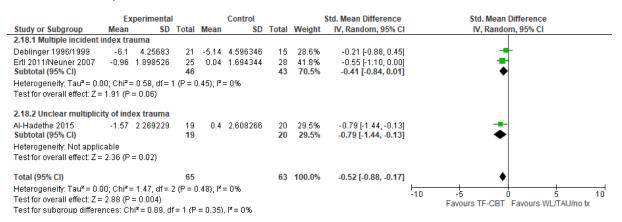


Figure 24: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 2 year follow-up (CDI change score)

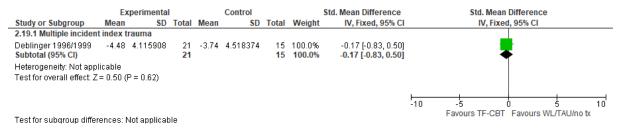


Figure 25: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems (SDQ-A change score)

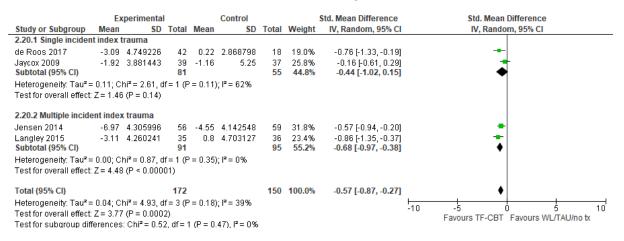


Figure 26: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Externalizing (CBCL Externalizing change score); Multiple incident index trauma

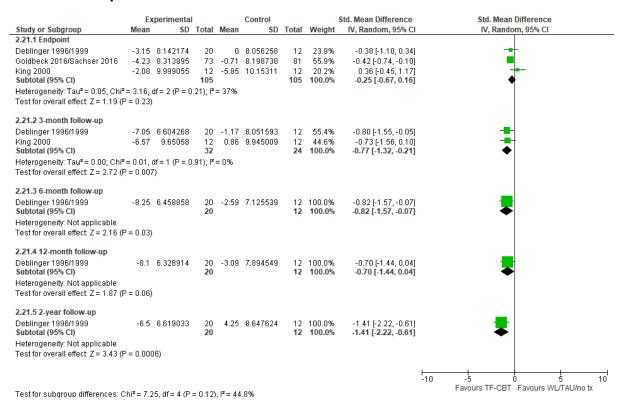


Figure 27: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Internalizing (CBCL Internalizing change score); Multiple incident index trauma

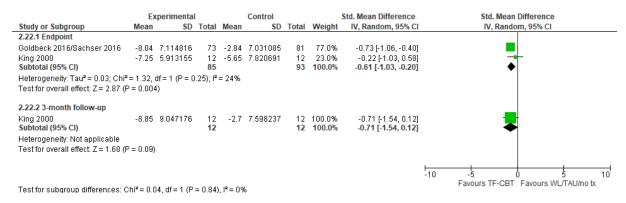


Figure 28: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Behaviour problems (CBCL total score; change score)

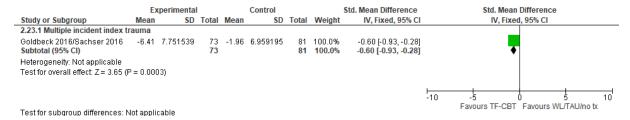


Figure 29: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Quality of life (KIDSCREEN-27: Global HRQoL T-scores/ILK; change score)

|   | E                    | cperimenta    | I         |            | Control  |       |        | Std. Mean Difference | Std. Mean Difference                |
|---|----------------------|---------------|-----------|------------|----------|-------|--------|----------------------|-------------------------------------|
| Study or Subgroup                           | Mean                 | SD            | Total     | Mean       | SD       | Total | Weight | IV, Random, 95% CI   | IV, Random, 95% CI                  |
| 2.24.1 Single incident index tra            | auma                 |               |           |            |          |       |        |                      |                                     |
| de Roos 2017                                | 7.48                 | 12.46031      | 42        | 1.07       | 11.14915 | 18    | 23.7%  | 0.52 [-0.04, 1.08]   | <del>-</del>                        |
| Subtotal (95% CI)                           |                      |               | 42        |            |          | 18    | 23.7%  | 0.52 [-0.04, 1.08]   | <b>◆</b>                            |
| Heterogeneity: Not applicable               |                      |               |           |            |          |       |        |                      |                                     |
| Test for overall effect: $Z = 1.83$ (       | P = 0.07             | )             |           |            |          |       |        |                      |                                     |
| 2.24.2 Multiple incident index t            | trauma               |               |           |            |          |       |        |                      |                                     |
| Goldbeck 2016/Sachser 2016                  | 7.07                 | 11.44655      |           | 3.91       | 11.74476 | 83    | 76.3%  | 0.27 [-0.04, 0.58]   | ·                                   |
| Subtotal (95% CI)                           |                      |               | 76        |            |          | 83    | 76.3%  | 0.27 [-0.04, 0.58]   | •                                   |
| Heterogeneity: Not applicable               |                      |               |           |            |          |       |        |                      |                                     |
| Test for overall effect: Z = 1.70 (         | P = 0.09             | )             |           |            |          |       |        |                      |                                     |
| Total (95% CI)                              |                      |               | 118       |            |          | 101   | 100.0% | 0.33 [0.06, 0.60]    | •                                   |
| Heterogeneity: Tau <sup>2</sup> = 0.00; Chi | <sup>2</sup> = 0.59. | df = 1 (P = 0 | ).44): I² | = 0%       |          |       |        |                      |                                     |
| Test for overall effect: Z = 2.38 (         |                      | ,             |           |            |          |       |        |                      | -10 -5 0 5 10                       |
| Test for subaroup differences: (            |                      |               | = 0.441   | 1  2 = 119 | 6        |       |        |                      | Favours WL/TAU/no tx Favours TF-CBT |

Figure 30: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Functional impairment at endpoint (CAPS: Functional impairment/SAS-SR-Y/Child Diagnostic Interview Schedule:Sum score of 7 areas of funcctional impairment; change score)

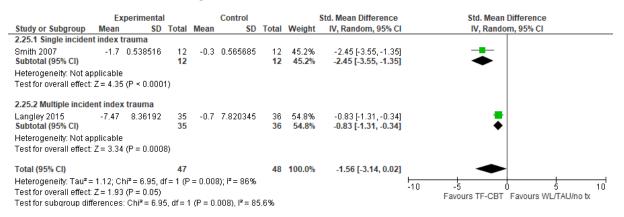


Figure 31: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Functional impairment at 3-month follow-up (CAPS: Functional impairment; change score)

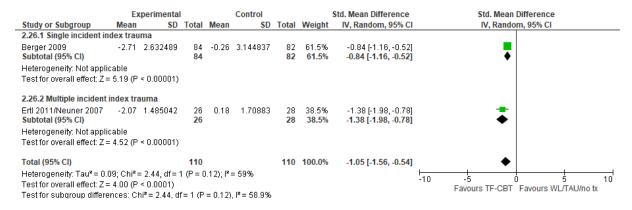


Figure 32: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Functional

## impairment at 6-12 month follow-up (CAPS: Functional impairment; change score); Multiple incident index trauma

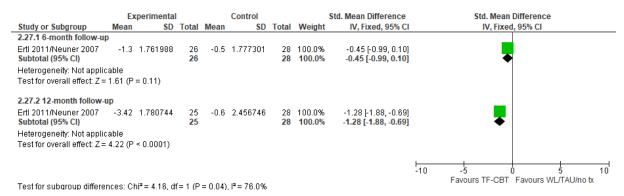


Figure 33: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning at endpoint (CGAS/fCPSS/GAF change score)

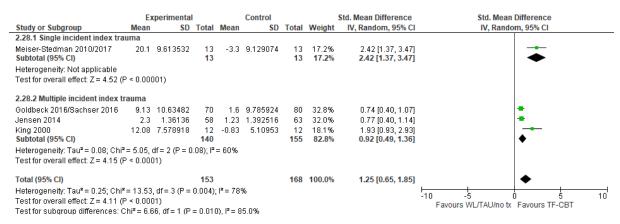


Figure 34: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning at 3-month follow-up (GAF; change score)

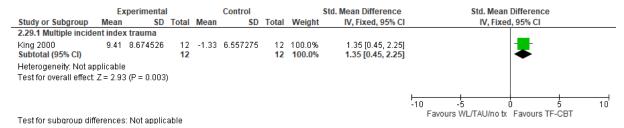
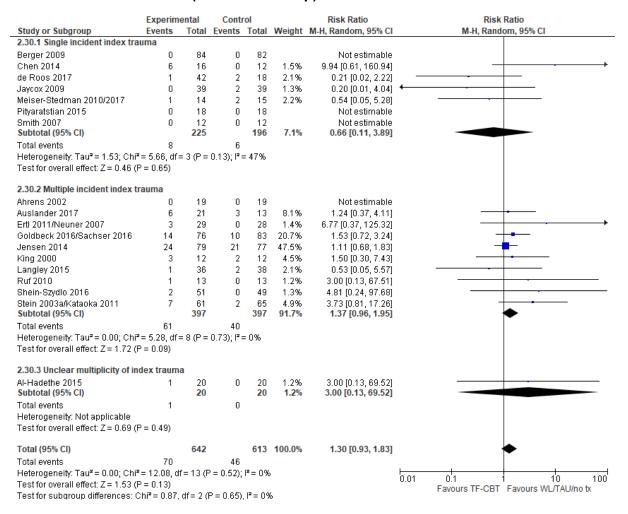


Figure 35: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Discontinuation (loss to follow-up)



Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 36: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

### symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA PTSD-RI/CPSS change score)

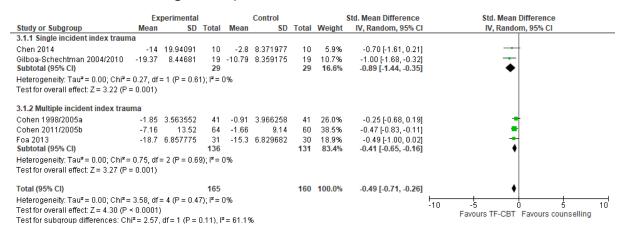


Figure 37: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 3-month follow-up (CRIES change score)

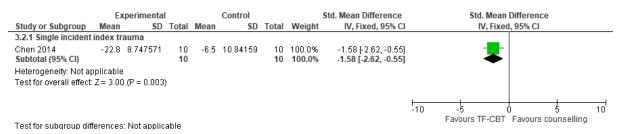


Figure 38: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 6-month follow-up (TSCC-PTSD/VCPSS change score)

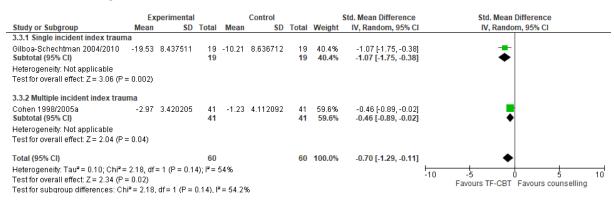


Figure 39: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

# symptomatology self-rated at 12-17 month follow-up (TSCC-PTSD/CPSS change score)

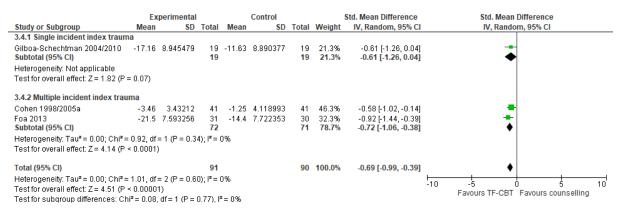


Figure 40: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

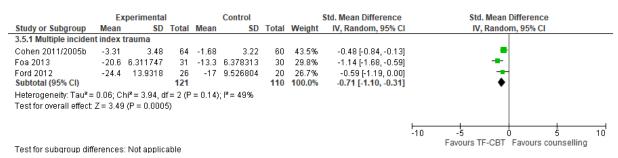


Figure 41: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at 3-month follow-up (CAPS change score)

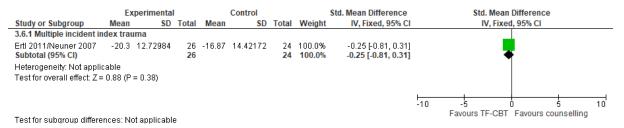


Figure 42: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at 6-month follow-up (CAPS change score)

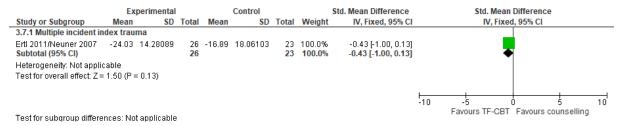


Figure 43: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at 12-month follow-up (CAPS/CPSS-I change score)

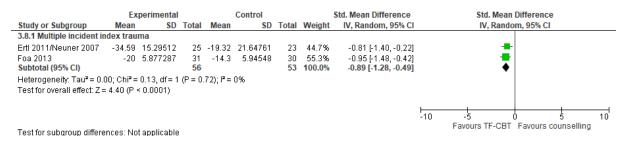


Figure 44: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at endpoint (number of people no longer meeting diagnostic criteria for PTSD)

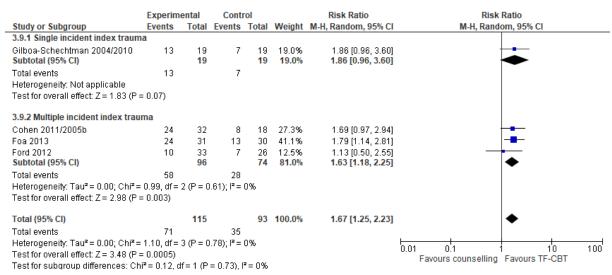


Figure 45: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at

### 6-month follow-up (number of people no longer meeting diagnostic criteria for PTSD)

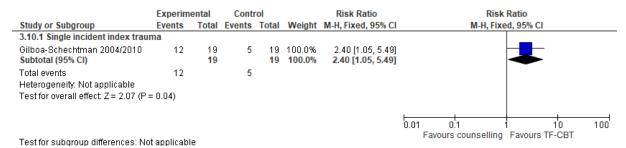


Figure 46: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at 12-month follow-up (number of people no longer meeting diagnostic criteria for PTSD)

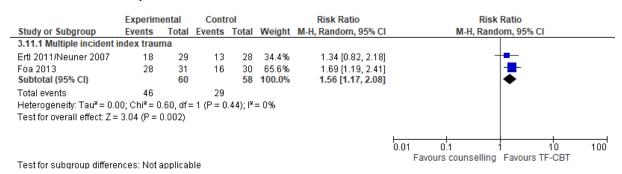


Figure 47: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Response at endpoint (number of people showing clinically significant improvement [based on RCI])

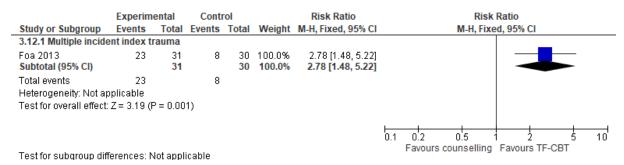


Figure 48: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Response at

## 12-month follow-up (number of people showing clinically significant improvement [based on RCI])

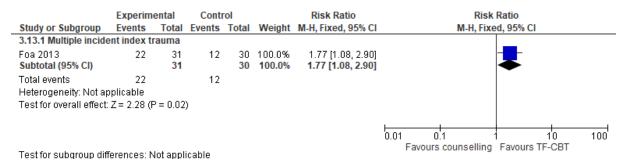


Figure 49: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Dissociative symptoms (TSCC-Dissociation change score); multiple incident index trauma

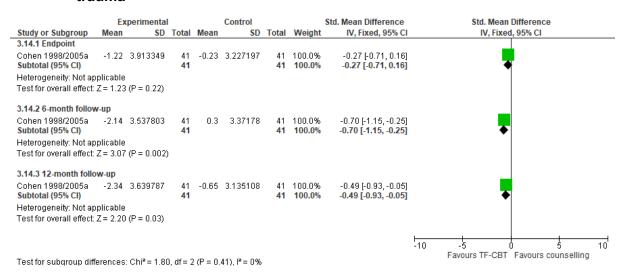


Figure 50: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety

### symptoms (STAI-State/SCARED/TSCC:Anxiety change score); Multiple incident index trauma

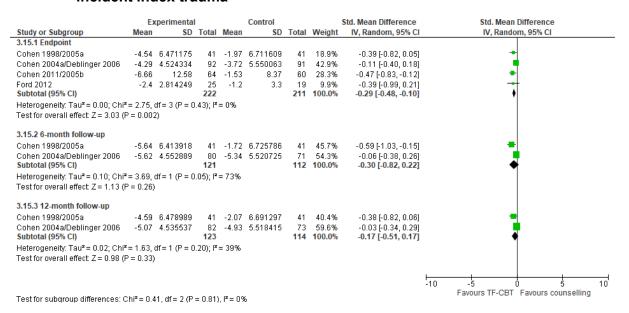


Figure 51: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at endpoint (BDI/CES-D/CDI/TSCC:Depression change score)

|  | Experimental Control |                |                      |        |          |       |        | Std. Mean Difference | Std. Mean Difference               |   |  |  |
|--|----------------------|----------------|----------------------|--------|----------|-------|--------|----------------------|------------------------------------|---|--|--|
| Study or Subgroup  | Mean                 | SD             | Total                | Mean   | SD       | Total | Weight | IV, Random, 95% CI   | I IV, Random, 95% CI               |   |  |  |
| 3.16.1 Single incident index traus                         | ma                   |                |                      |        |          |       |        |                      |                                    |   |  |  |
| Chen 2014  | -3.7                 | 9.681942       | 10                   | 3.9    | 7.107039 | 10    | 6.2%   | -0.86 [-1.78, 0.07]  | ı <del></del>                      |   |  |  |
| Gilboa-Schechtman 2004/2010                                | -13.95               | 8.747565       | 19                   | -6.94  | 6.867117 | 19    | 10.0%  | -0.87 [-1.54, -0.20] | <u></u>                            |   |  |  |
| Subtotal (95% CI)  |                      |                | 29                   |        |          | 29    | 16.2%  | -0.87 [-1.41, -0.33] | •                                  |   |  |  |
| Heterogeneity: Tau² = 0.00; Chi² =                         | 0.00, df             | = 1 (P = 0.9)  | $ 8\rangle;  ^2 = 1$ | 0%     |          |       |        |                      |                                    |   |  |  |
| Test for overall effect: Z = 3.14 (P =                     | = 0.002)             |                |                      |        |          |       |        |                      |                                    |   |  |  |
| 3.16.2 Multiple incident index tra                         | uma                  |                |                      |        |          |       |        |                      |                                    |   |  |  |
| Cohen 1998/2005a   | -4.76                | 5.861467       | 41                   | -0.25  | 5.476148 | 41    | 16.0%  | -0.79 [-1.24, -0.34] | ] <del>-</del>                     |   |  |  |
| Cohen 2004a/Deblinger 2006                                 | -4.22                | 4.963205       | 92                   | -3.32  | 6.3916   | 91    | 22.5%  | -0.16 [-0.45, 0.13]  | i <del>*</del>                     |   |  |  |
| Cohen 2011/2005b   | -2.44                | 6.02           | 64                   | -1.03  | 3.89     | 60    | 19.7%  | -0.27 [-0.63, 0.08]  | i <del>-</del>                     |   |  |  |
| Foa 2013   | -11.2                | 5.41147        | 31                   | -8.4   | 5.467092 | 30    | 14.0%  | -0.51 [-1.02, 0.00]  | ] <del></del>                      |   |  |  |
| Ford 2012  | -2.3                 | 2.525866       | 25                   | -2.6   | 2.834608 | 19    | 11.6%  | 0.11 [-0.49, 0.71]   | <u>+</u>                           |   |  |  |
| Subtotal (95% CI)  |                      |                | 253                  |        |          | 241   | 83.8%  | -0.33 [-0.59, -0.06] | 1 ♦                                |   |  |  |
| Heterogeneity: Tau² = 0.04; Chi² =                         | 7.92, df             | = 4 (P = 0.0)  | $9); I^2 = -$        | 49%    |          |       |        |                      |                                    |   |  |  |
| Test for overall effect: Z = 2.42 (P =                     | = 0.02)              |                |                      |        |          |       |        |                      |                                    |   |  |  |
| Total (95% CI)   |                      |                | 282                  |        |          | 270   | 100.0% | -0.41 [-0.67, -0.16] | ı •                                |   |  |  |
| Heterogeneity: Tau <sup>2</sup> = 0.05; Chi <sup>2</sup> = | 11.65. d             | lf = 6 (P = 0. | 07): <b> ²</b> =     | 49%    |          |       |        | - / -                | ·                                  | 4 |  |  |
| Test for overall effect: Z = 3.17 (P =                     |                      |                | 71 -                 |        |          |       |        |                      | -10 -5 0 5 10                      | J |  |  |
| Test for subgroup differences: Ch                          |                      | df = 1 (P =    | 0.08). P             | e 67.8 | %        |       |        |                      | Favours TF-CBT Favours counselling |   |  |  |
|  |                      |                |                      |        |          |       |        |                      |                                    |   |  |  |

Figure 52: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 3-month follow-up (CES-D/MINI:Depression change score)

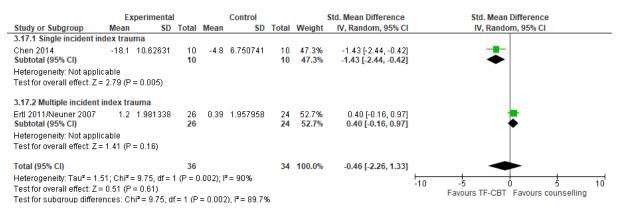


Figure 53: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 6-month follow-up (BDI/CDI/MINI:Depression change score)

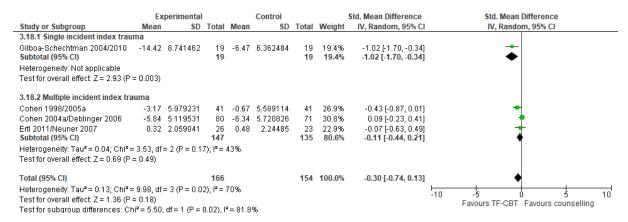


Figure 54: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression

### symptoms at 12-17 month follow-up (BDI/CDI/MINI:Depression change score)

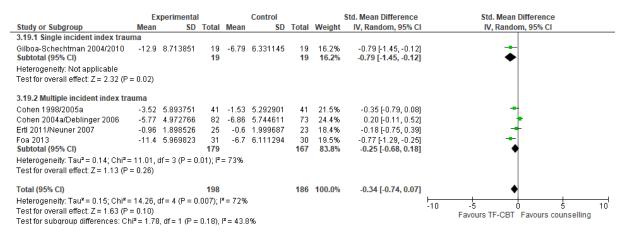


Figure 55: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Internalizing (CBCL Internalizing change score); Multiple incident index trauma

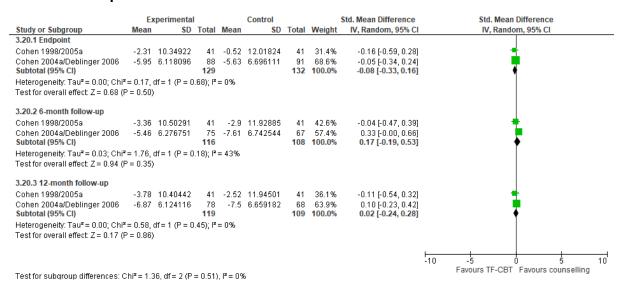


Figure 56: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional

Test for subgroup differences:  $Chi^2 = 1.36$ , df = 2 (P = 0.51),  $I^2 = 0\%$ 

#### and behavioural problems-Externalizing (CBCL Externalizing change score); Multiple incident index trauma

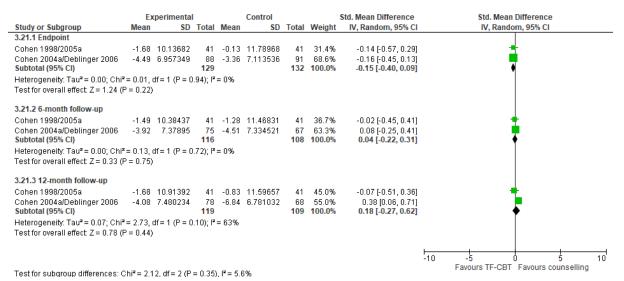


Figure 57: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Behaviour

problems (CBCL total score; change score); multiple incident index trauma

Experimental Control Std. Mean Difference Std. Mean Difference SD Total Weight Study or Subgroup Mean SD Total Mean IV, Random, 95% CI IV. Random, 95% CI 3.22.1 Endpoint Cohen 1998/2005a -2.68 10.62618 41 -0.47 -0.19 [-0.63, 0.24] Cohen 2004a/Deblinger 2006 -17.03 18.47255 88 -13.5 19.50769 91 46.5% -0.18 [-0.48, 0.11] 20.45 Cohen 2011/2005b -8.78 19.98 64 -10.12 60 32.3% 0.07 (-0.29 0.42) -0.11 [-0.31, 0.09] Subtotal (95% CI) Heterogeneity:  $Tau^2 = 0.00$ ;  $Chi^2 = 1.35$ , df = 2 (P = 0.51);  $I^2 = 0\%$ Test for overall effect: Z = 1.04 (P = 0.30) 3.22.2 6-month follow-up 
 Cohen 1998/2005a
 -3.22
 10.87942
 41
 -2.12
 11.72829

 Cohen 2004a/Deblinger 2006
 -16.04
 19.44402
 75
 -19.71
 20.59929

 Subtotal (95% CI)
 116
 41 36.8% -0.10 [-0.53, 0.34] 63.2% 0.18 (-0.15, 0.51) 0.08 [-0.18, 0.34] Heterogeneity:  $Tau^2 = 0.00$ ;  $Chi^2 = 1.01$ , df = 1 (P = 0.32);  $I^2 = 1\%$ Test for overall effect: Z = 0.59 (P = 0.55) 3.22.3 12-month follow-up 
 Cohen 1998/2005a
 -3.59
 11.1094
 41
 -1.57
 11.85405

 Cohen 2004a/Deblinger 2006
 -19.31
 18.83627
 78
 -23.16
 18.93715
 41 42.5% -0.17 [-0.61, 0.26] 68 57.5% 0.20 f-0.12, 0.531 119 109 100.0% 0.04 [-0.32, 0.41] Heterogeneity:  $Tau^2 = 0.03$ ;  $Chi^2 = 1.85$ , df = 1 (P = 0.17);  $I^2 = 46\%$ Test for overall effect: Z = 0.23 (P = 0.82) -10 10 Favours TF-CBT Favours counselling

Figure 58: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Functional

### impairment (CAPS: Functional impairment; change score); Multiple incident index trauma

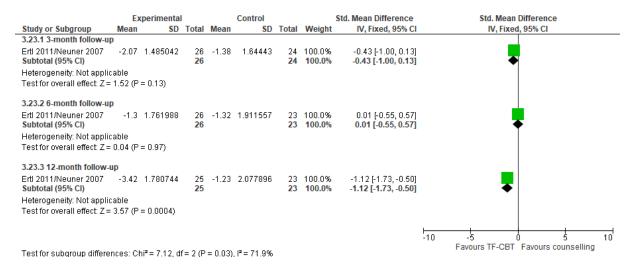


Figure 59: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning at endpoint (CGAS; change score)

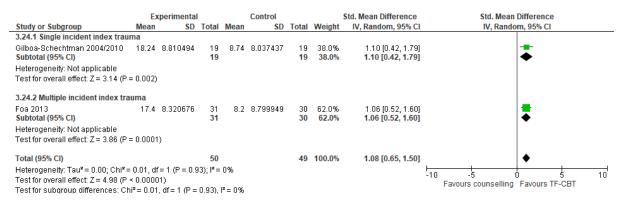


Figure 60: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning at 6-month follow-up (CGAS; change score)

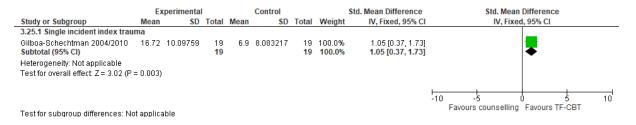


Figure 61: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning at 12-month follow-up (CGAS; change score)

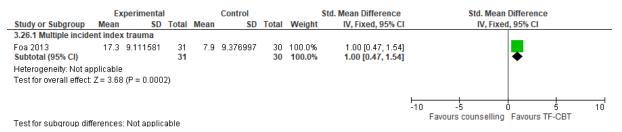
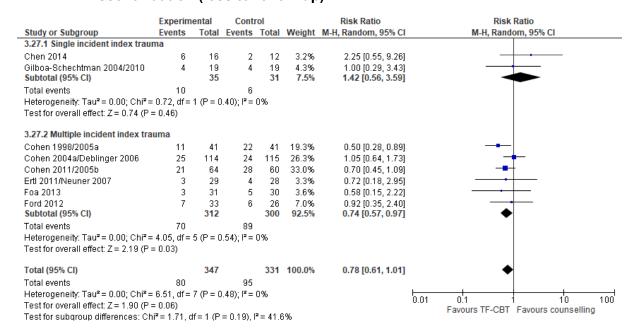


Figure 62: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Discontinuation (loss to follow-up)



Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically important symptoms/PTSD - Single incident index trauma

Figure 63: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically

## important symptoms/PTSD: PTSD symptomatology self-rated (CRTI change score); Single incident index trauma

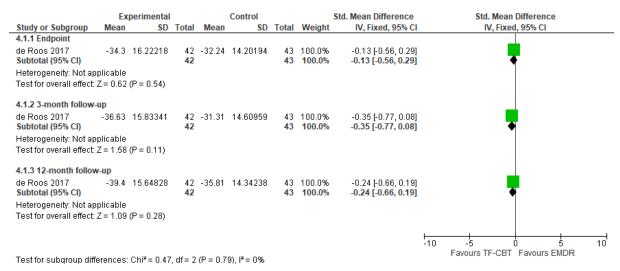


Figure 64: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (CAPS-CA change score)

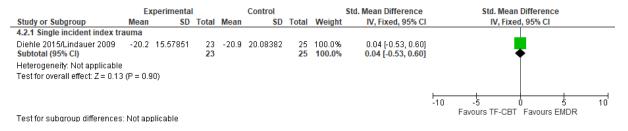


Figure 65: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically

#### important symptoms/PTSD: Emotional and behavioural problems (SDQ-A change score); Single incident index trauma

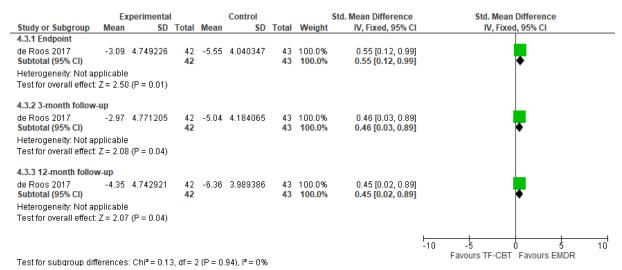
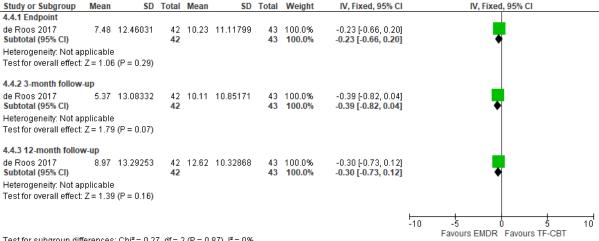


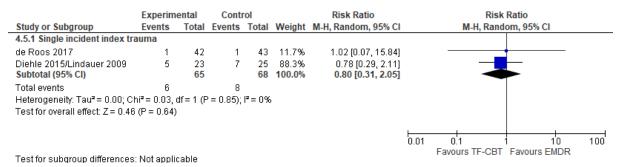
Figure 66: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Quality of life (KIDSCREEN-27: Global HRQoL T-

scores; change score); Single incident index trauma Std. Mean Difference Experimental Control Std. Mean Difference Mean SD Total Mean SD Total Weight IV, Fixed, 95% CI IV, Fixed, 95% CI



Test for subgroup differences:  $Chi^2 = 0.27$ , df = 2 (P = 0.87),  $I^2 = 0\%$ 

Figure 67: Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 68: Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated (SPTSS change score); Unclear multiplicity of trauma

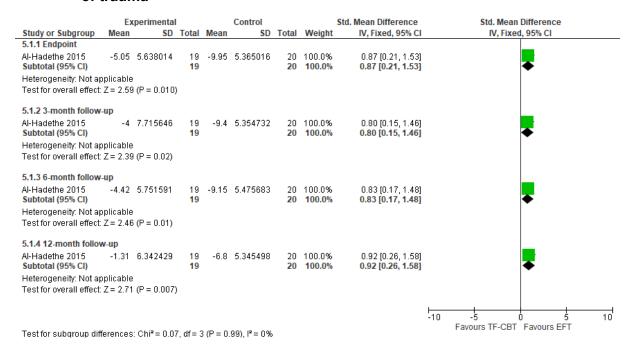


Figure 69: Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

#### Anxiety symptoms (HADS-A change score); Unclear multiplicity of index trauma

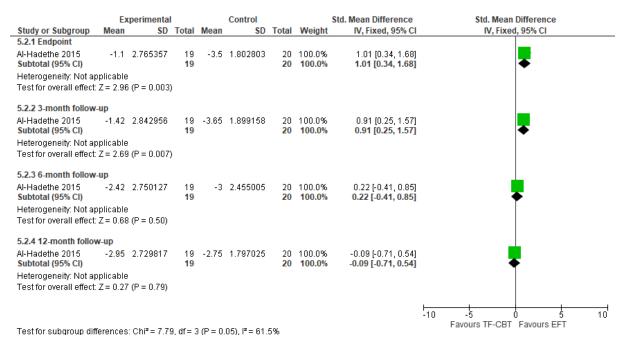


Figure 70: Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Depression symptoms (HADS-D change score); Unclear multiplicity of index trauma

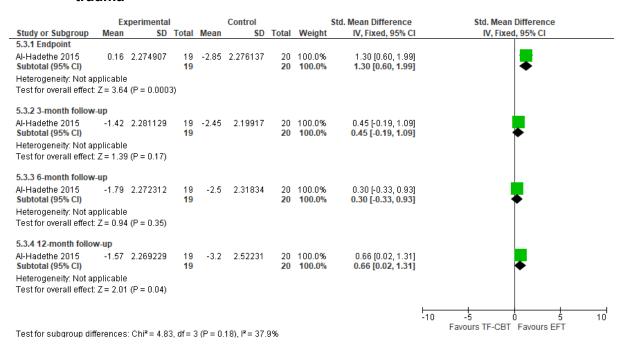
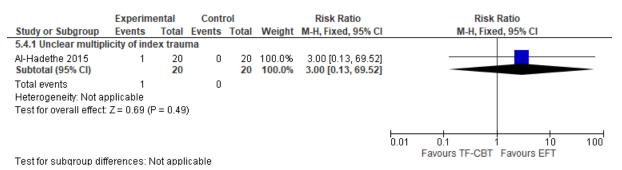


Figure 71: Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 72: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (ADIS-C: PTSD; change score); Multiple incident index trauma

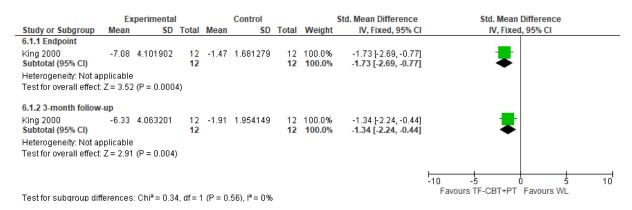


Figure 73: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms (RCMAS; change score); Multiple incident index trauma

|   | Ex      | perimental   | ı               |         | Control      |                 |        | Std. Mean Difference | Std. Mean Difference         |
|---|---------|--------------|-----------------|---------|--------------|-----------------|--------|----------------------|------------------------------|
| Study or Subgroup                                 | Mean    | \$D          | Total           | Mean    | SD           | Total           | Weight | IV, Fixed, 95% CI    | IV, Fixed, 95% CI            |
| 6.2.1 Endpoint                                    |         |              |                 |         |              |                 |        |                      |                              |
| King 2000<br>Subtotal (95% CI)                    | -5.75   | 14.15629     | 12<br><b>12</b> | -1.59   | 10.11676     | 12<br><b>12</b> |        |                      | -                            |
| Heterogeneity: Not ap<br>Test for overall effect: |         |              |                 |         |              |                 |        |                      |                              |
| 6.2.2 3-month follow-                             | -up     |              |                 |         |              |                 |        |                      |                              |
| King 2000<br>Subtotal (95% CI)                    | -9.59   | 10.82639     | 12<br><b>12</b> | -1.59   | 9.807839     | 12<br><b>12</b> |        |                      | -                            |
| Heterogeneity: Not ap<br>Test for overall effect: |         |              |                 |         |              |                 |        |                      |                              |
|   |         |              |                 |         |              |                 |        |                      | -10 -5 0 5 10                |
| Test for subgroup diff                            | erences | : Chi² = 0.5 | 1, df = 1       | (P = 0. | 48), I² = 0% |                 |        |                      | Favours TF-CBT+PT Favours WL |

Figure 74: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms (CDI; change score); Multiple incident index trauma

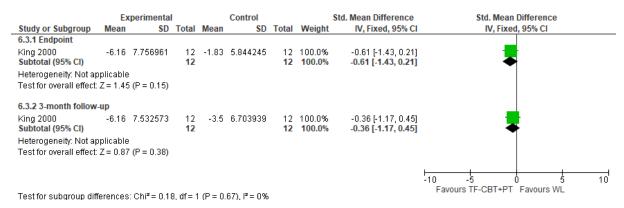


Figure 75: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Internalizing (CBCL: Internalizing; change score); Multiple incident index trauma

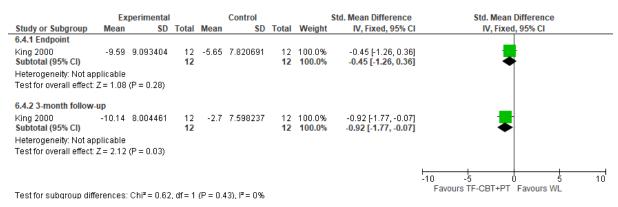


Figure 76: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Externalizing (CBCL: Externalizing; change score); Multiple incident index trauma

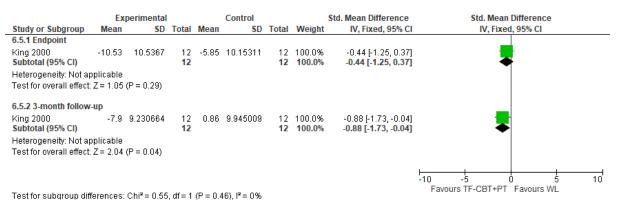


Figure 77: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning (GAF; change score); Multiple incident index trauma

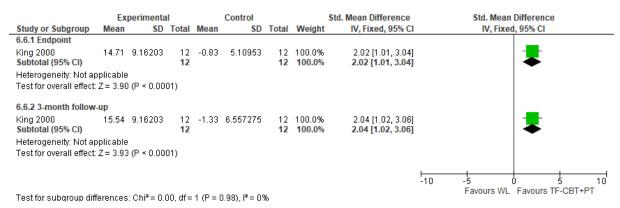
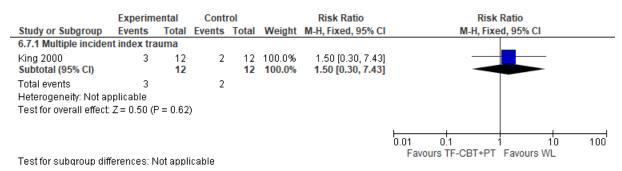


Figure 78: Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Discontinuation (loss to follow-up)



Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 79: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (ADIS-C: PTSD; change score); Multiple incident index trauma

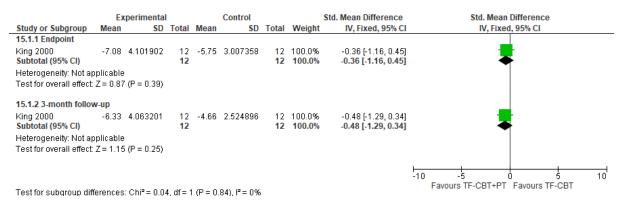


Figure 80: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms (RCMAS; change score); Multiple incident index trauma

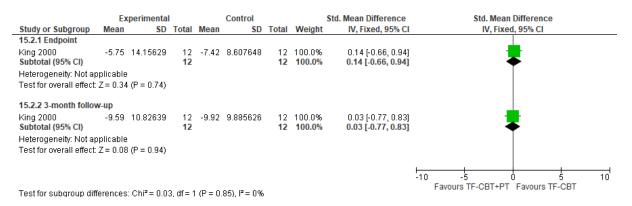


Figure 81: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms (CDI; change score); Multiple incident index trauma

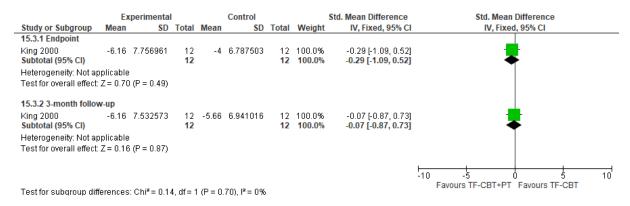


Figure 82: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important

## symptoms/PTSD: Emotional and behavioural problems-Internalizing (CBCL: Internalizing; change score); Multiple incident index trauma

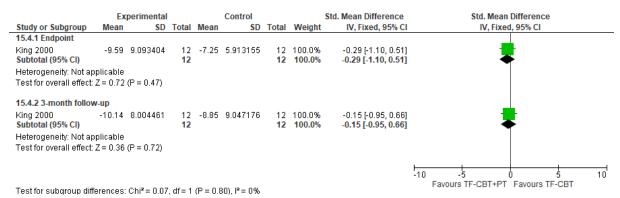


Figure 83: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Externalizing (CBCL: Externalizing; change score); Multiple incident index trauma

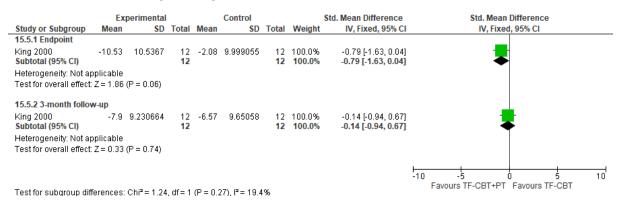


Figure 84: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning (GAF; change score); Multiple incident index trauma

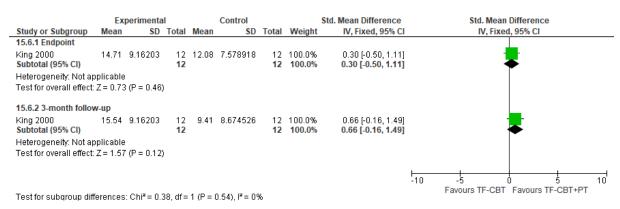
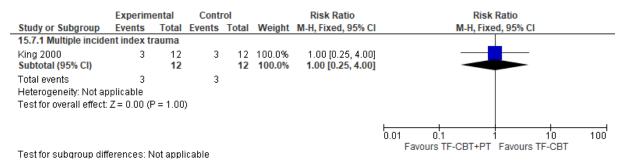


Figure 85: Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Trauma-focused CBT versus parent training (CBT with parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 86: Trauma-focused CBT versus parent training (CBT with parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (K-SADS-E: PTSD; change score); Multiple incident index trauma

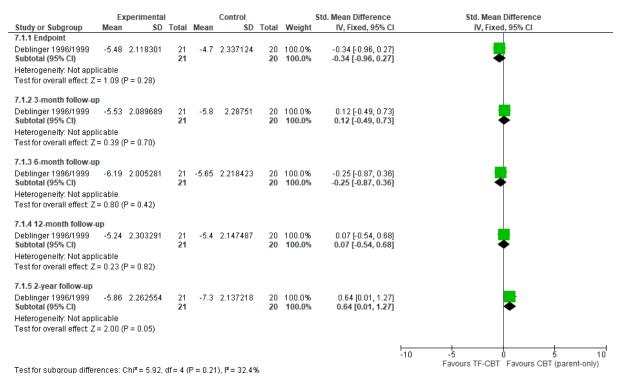
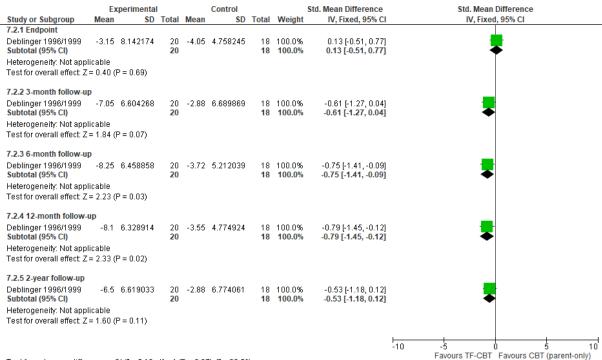


Figure 87: Trauma-focused CBT versus parent training (CBT with parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

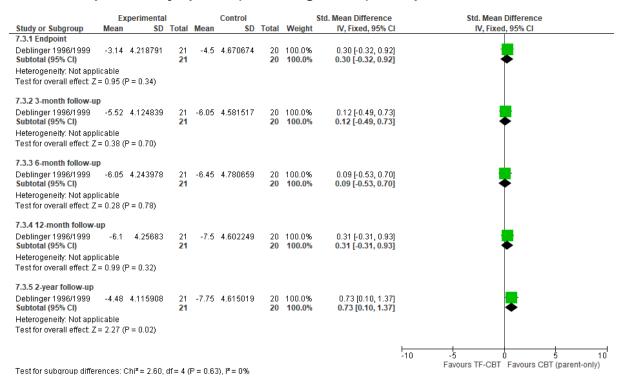
## Emotional and behavioural problems-Externalizing (CBCL Externalizing change score); Multiple incident index trauma



Test for subgroup differences: Chi<sup>2</sup> = 5.16, df = 4 (P = 0.27), I<sup>2</sup> = 22.5%

Figure 88: Trauma-focused CBT versus parent training (CBT with parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Depression symptoms (CDI change score); Multiple incident index trauma



Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 89: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

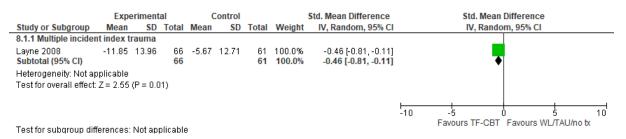


Figure 90: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically

#### important symptoms/PTSD: PTSD symptomatology self-rated at 4 month follow-up (IES/SPTSS/CRIES/UCLA PTSD-RI/CPTS-RI change score)

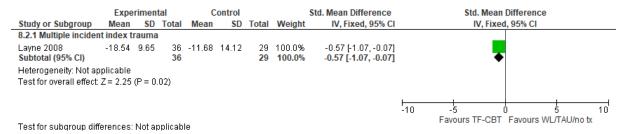


Figure 91: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Response at endpoint (number of people showing clinically significant improvement, based on reliable change indices [RCI]/rated as 'much/very much improved' on CGI)

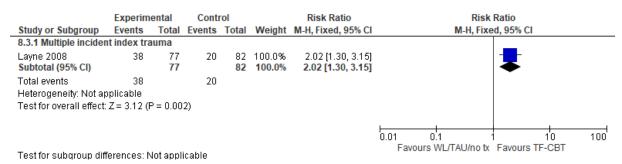


Figure 92: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Response at 4-month follow-up (number of people showing clinically significant improvement, based on reliable change indices [RCI])

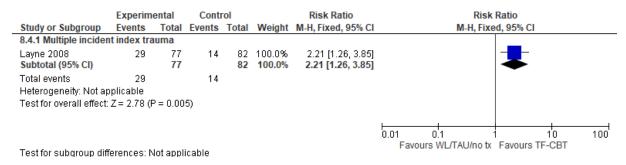


Figure 93: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically

## important symptoms/PTSD: Depression symptoms at endpoint (HADS-D/CES-D/CDI/MFQ/DSRS/BDI change score)

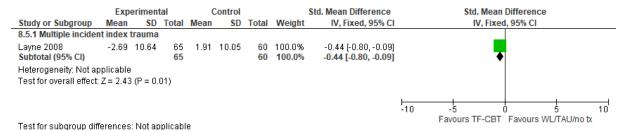


Figure 94: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 4 month follow-up (BDI/HADS-D/CES-D/CDI/MINI:Depression /DSRS change score)

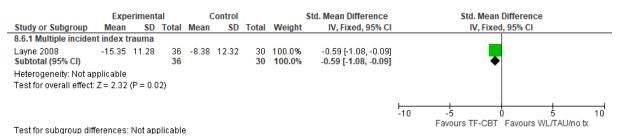
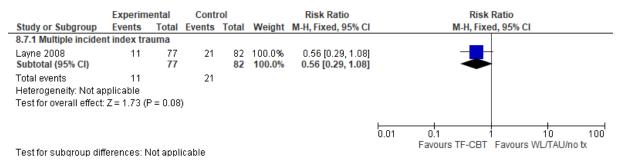


Figure 95: Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by specific intervention: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 96: Sub-analysis by specific intervention: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically

## important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

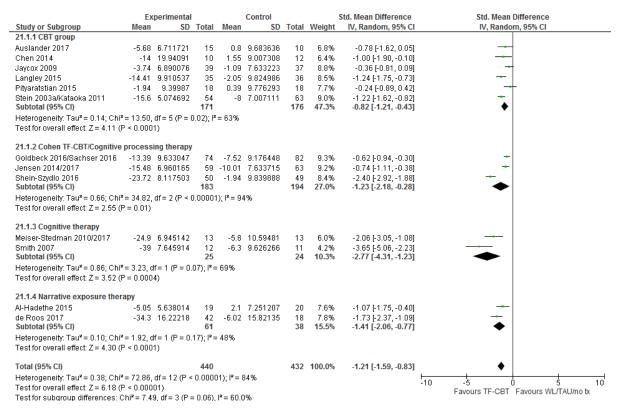
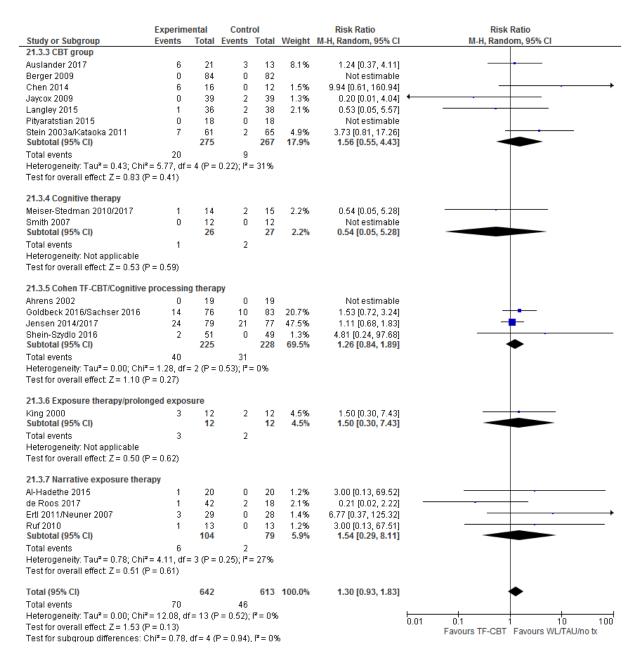


Figure 97: Sub-analysis by specific intervention: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically

## important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score)

|   | Ex        | perimental    |           |           | Control  |       |        | Std. Mean Difference | Std. Mean Difference |
|---|-----------|---------------|-----------|-----------|----------|-------|--------|----------------------|----------------------|
| Study or Subgroup   | Mean      |               |           | Mean      | SD       | Total | Weight | IV, Random, 95% CI   | IV, Random, 95% CI   |
| 21.2.1 Cohen TF-CBT/Cognitive   | process   | ing therapy   |           |           |          |       |        |                      |                      |
| Goldbeck 2016/Sachser 2016  | -26.35    | 17.33886      | 76        | -14.1     | 16.91013 | 83    | 19.0%  | -0.71 [-1.03, -0.39] | -                    |
| Jensen 2014/2017  | -29.64    | 16.75992      |           | -18.6     | 17.62647 | 61    | 18.6%  | -0.64 [-1.01, -0.26] | <u>†</u>             |
|   |           |               |           |           |          | 144   | 37.6%  | -0.68 [-0.92, -0.44] | •                    |
|   |           |               | 76); l² = | : 0%      |          |       |        |                      |                      |
| Test for overall effect: $Z = 5.47$ (F  | o < 0.000 | 01)           |           |           |          |       |        |                      |                      |
|   |           |               |           |           |          |       |        |                      |                      |
|   |           |               |           |           |          |       |        |                      |                      |
|   |           |               |           |           |          |       |        |                      | <del></del>          |
|   | -48.9     | 12.01499      |           | -14.4     | 12.1359  |       |        |                      | <del></del>          |
|   |           |               |           |           |          | 25    | 21.8%  | -2.80 [-3.62, -1.99] | •                    |
|   |           |               | 92); l² = | = 0%      |          |       |        |                      |                      |
| Lest for overall effect: ∠= 6.73 (F   | ° < 0.000 | U1)           |           |           |          |       |        |                      |                      |
| 21.2.3 Exposure therapy/prolon  | nned eyn  | nsure         |           |           |          |       |        |                      |                      |
|   |           |               | 21        | 2.20      | 2 220510 | 1.4   | 15 200 | 0.07 ( 4.60 . 0.25)  |                      |
|   |           |               |           |           |          |       |        |                      |                      |
|   | -3.73     | 3.007330      | 33        | -1.47     | 1.001273 |       |        |                      | •                    |
| * *   | = 1.42 d  | f = 1 (P = 0  | 23): 12 = | 30%       |          |       |        |                      | •                    |
|   |           |               | 20),1 -   | - 30 %    |          |       |        |                      |                      |
| 1001101 0401011 011002 Z = 0.04 (I  | - 0.000   | 7/            |           |           |          |       |        |                      |                      |
| 21.2.4 Narrative exposure there   | ару       |               |           |           |          |       |        |                      |                      |
| Ruf 2010  | -26.1     | 9.750897      | 12        | -4.5      | 12.33937 | 13    | 12.7%  | -1.87 [-2.84, -0.90] | <del></del>          |
| Subtotal (95% CI)   |           |               | 12        |           |          | 13    | 12.7%  | -1.87 [-2.84, -0.90] | <b>◆</b>             |
| Heterogeneity: Not applicable   |           |               |           |           |          |       |        |                      |                      |
| Test for overall effect: Z = 3.79 (F  | P = 0.000 | 2)            |           |           |          |       |        |                      |                      |
|   |           |               |           |           |          |       |        |                      |                      |
| Study or Subgroup   Mean   SD   Total   Mean   SD   SD   SD   SD   SD   SD   SD   S |           |               |           |           |          |       |        |                      |                      |
|   |           | ,             | 0.0001)   | ; I² = 80 | %        |       |        |                      | -10 -5 0 5 10        |
| •   |           |               |           |           |          |       |        |                      |                      |
| Test for subgroup differences: C  | hi² = 28. | 74, df = 3 (P | < 0.00    | 001), I²  | = 89.6%  |       |        |                      |                      |

Figure 98: Sub-analysis by specific intervention: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by format: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 99: Sub-analysis by format: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important

## symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change

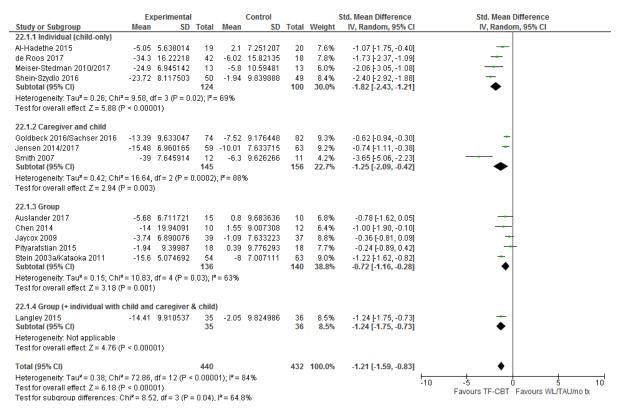
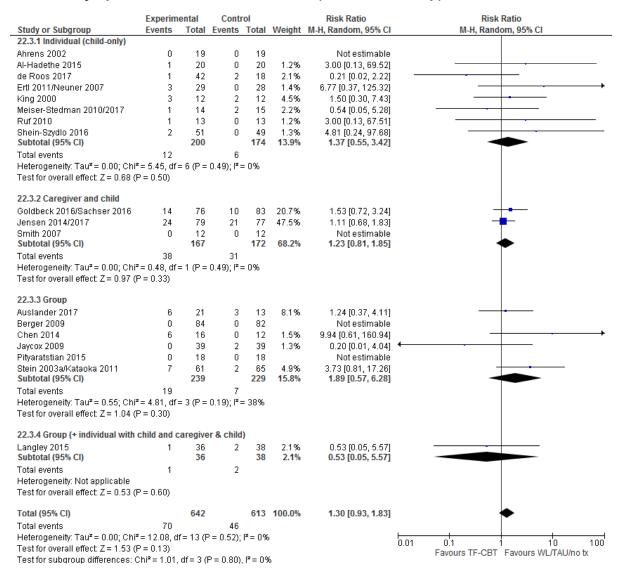


Figure 100: Sub-analysis by format: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C: PTSD/CPTSDI; change score)

|  | Ex            | perimental     |           |                      | Control  |       |        | Std. Mean Difference |     | Std. Mean Difference                    |      |
|--|---------------|----------------|-----------|----------------------|----------|-------|--------|----------------------|-----|---|------|
| Study or Subgroup  | Mean          | SD             | Total     | Mean                 | SD       | Total | Weight | IV, Random, 95% CI   |     | IV, Random, 95% CI                      |      |
| 22.2.1 Individual (child-only)                             |               |                |           |                      |          |       |        |                      |     |   |      |
| Deblinger 1996/1999  | -5.48         | 2.118301       | 21        | -3.29                | 2.339519 | 14    | 15.2%  | -0.97 [-1.69, -0.25] |     |   |      |
| King 2000  | -5.75         | 3.007358       | 12        | -1.47                | 1.681279 | 12    | 12.7%  | -1.70 [-2.65, -0.74] |     | <del></del>                             |      |
| Meiser-Stedman 2010/2017                                   | -9.3          | 2.433105       | 13        | -1.5                 | 2.862691 | 13    | 11.1%  | -2.84 [-3.98, -1.71] |     | <del></del>                             |      |
| Ruf 2010   | -26.1         | 9.750897       | 12        | -4.5                 | 12.33937 | 13    | 12.7%  | -1.87 [-2.84, -0.90] |     | <del></del>                             |      |
| Subtotal (95% CI)  |               |                | 58        |                      |          | 52    | 51.7%  | -1.76 [-2.52, -1.01] |     | <b>◆</b>                                |      |
| Heterogeneity: Tau <sup>2</sup> = 0.36; Chi <sup>2</sup> : | = 7.88, d     | f = 3 (P = 0.  | 05); l² = | 62%                  |          |       |        |                      |     |   |      |
| Test for overall effect: Z = 4.59 (P                       | < 0.000       | 01)            |           |                      |          |       |        |                      |     |   |      |
|  |               |                |           |                      |          |       |        |                      |     |   |      |
| 22.2.2 Caregiver and child                                 |               |                |           |                      |          |       |        |                      |     |   |      |
| Goldbeck 2016/Sachser 2016                                 | -26.35        | 17.33886       | 76        | -14.1                | 16.91013 | 83    | 19.0%  | -0.71 [-1.03, -0.39] |     | -                                       |      |
| Jensen 2014/2017   | -29.64        | 16.75992       | 55        | -18.6                | 17.62647 | 61    | 18.6%  | -0.64 [-1.01, -0.26] |     | *                                       |      |
| Smith 2007   | -48.9         | 12.01499       | 12        | -14.4                | 12.1359  | 12    | 10.8%  | -2.76 [-3.93, -1.59] |     |   |      |
| Subtotal (95% CI)  |               |                | 143       |                      |          | 156   | 48.3%  | -1.08 [-1.77, -0.40] |     | <b>◆</b>                                |      |
| Heterogeneity: Tau <sup>2</sup> = 0.27; Chi <sup>2</sup> : | = 11.71.      | df = 2 (P = I  | 0.003);   | I <sup>2</sup> = 839 | 5        |       |        |                      |     |   |      |
| Test for overall effect: Z = 3.09 (P                       |               |                |           |                      |          |       |        |                      |     |   |      |
| •  |               |                |           |                      |          |       |        |                      |     |   |      |
| Total (95% CI)   |               |                | 201       |                      |          | 208   | 100.0% | -1.47 [-2.03, -0.90] |     | <b>♦</b>                                |      |
| Heterogeneity: Tau <sup>2</sup> = 0.41; Chi <sup>2</sup> : | = 30.42,      | df = 6 (P < 1  | 0.0001)   | $     ^2 = 80$       | %        |       |        |                      | 10  | <del></del>                             |      |
| Test for overall effect: Z = 5.11 (P                       | < 0.000       | 01)            |           |                      |          |       |        |                      | -10 | -5 Ó 5 Favours TF-CBT Favours WL/TAU/no | 10   |
| Test for subgroup differences: Cl                          | $hi^2 = 1.73$ | 3, df = 1 (P = | = 0.19),  | $I^2 = 42$ .         | 2%       |       |        |                      |     | PAVOUIS IF-CBT PAVOUIS WE/TAO/III       | ) LX |

Figure 101: Sub-analysis by format: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by age range: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 102: Sub-analysis by age range: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important

## symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

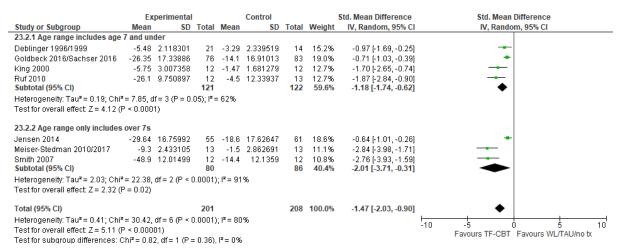
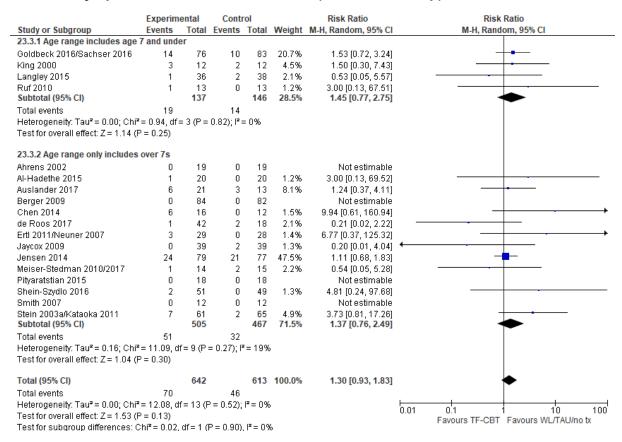


Figure 103: Sub-analysis by age range: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score)

|                                      | Experime                 | ntal           |               | Control  |       |        | Std. Mean Difference | Std. Mean Difference                |
|--------------------------------------|--------------------------|----------------|---------------|----------|-------|--------|----------------------|-------------------------------------|
| Study or Subgroup                    | Mean SD Total            |                |               | SD       | Total | Weight | IV, Random, 95% CI   | IV, Random, 95% CI                  |
| 23.2.1 Age range includes age        | 7 and under              |                |               |          |       |        |                      |                                     |
| Deblinger 1996/1999                  | -5.48 2.118              | 301 21         | -3.29         | 2.339519 | 14    | 15.2%  | -0.97 [-1.69, -0.25] |                                     |
| Goldbeck 2016/Sachser 2016           | -26.35 17.33             | 386 76         | -14.1         | 16.91013 | 83    | 19.0%  | -0.71 [-1.03, -0.39] | *                                   |
| King 2000                            | -5.75 3.007              | 358 12         | -1.47         | 1.681279 | 12    | 12.7%  | -1.70 [-2.65, -0.74] | <del></del>                         |
| Ruf 2010                             | -26.1 9.750              | 397 12         | -4.5          | 12.33937 | 13    | 12.7%  | -1.87 [-2.84, -0.90] | <del></del>                         |
| Subtotal (95% CI)                    |                          | 121            |               |          | 122   | 59.6%  | -1.18 [-1.74, -0.62] | <b>◆</b>                            |
| Heterogeneity: Tau² = 0.19; Chi²     | = 7.85, df = 3 (F        | $= 0.05); I^2$ | = 62%         |          |       |        |                      |                                     |
| Test for overall effect: Z = 4.12 (F | P < 0.0001)              |                |               |          |       |        |                      |                                     |
|                                      |                          |                |               |          |       |        |                      |                                     |
| 23.2.2 Age range only includes       | over 7s                  |                |               |          |       |        |                      |                                     |
| Jensen 2014                          | -29.64 16.75             | 992 55         | -18.6         | 17.62647 | 61    | 18.6%  | -0.64 [-1.01, -0.26] | *                                   |
| Meiser-Stedman 2010/2017             | -9.3 2.433               | 105 13         | -1.5          | 2.862691 | 13    | 11.1%  | -2.84 [-3.98, -1.71] | <del></del>                         |
| Smith 2007                           | -48.9 12.01              | 499 12         | -14.4         | 12.1359  | 12    | 10.8%  | -2.76 [-3.93, -1.59] | <del></del>                         |
| Subtotal (95% CI)                    |                          | 80             |               |          | 86    | 40.4%  | -2.01 [-3.71, -0.31] | •                                   |
| Heterogeneity: Tau² = 2.03; Chi²     | = 22.38, df = 2          | P < 0.0001     | $); I^2 = 91$ | %        |       |        |                      |                                     |
| Test for overall effect: Z = 2.32 (F | P = 0.02)                |                |               |          |       |        |                      |                                     |
|                                      |                          |                |               |          |       |        |                      |                                     |
| Total (95% CI)                       |                          | 201            |               |          | 208   | 100.0% | -1.47 [-2.03, -0.90] | <b>◆</b>                            |
| Heterogeneity: Tau² = 0.41; Chi²     | = 30.42, df = 6          | P < 0.0001     | $); I^2 = 80$ | 1%       |       |        |                      | -10 -5 0 5 10                       |
| Test for overall effect: Z = 5.11 (F | P < 0.00001)             |                |               |          |       |        |                      | Favours TF-CBT Favours WL/TAU/no tx |
| Test for subgroup differences: C     | $hi^2 = 0.82, df = 0.82$ | (P = 0.36)     | $I^2 = 0\%$   |          |       |        |                      | Tavodis II -ODT Tavodis WDTAO/IIO K |

Figure 104: Sub-analysis by age range: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 105: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months)

## of clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

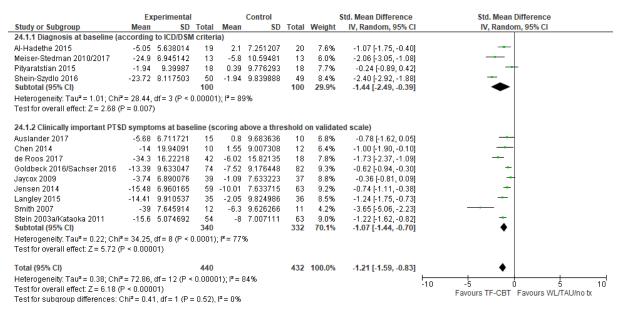


Figure 106: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C: PTSD/CPTSDI; change score)

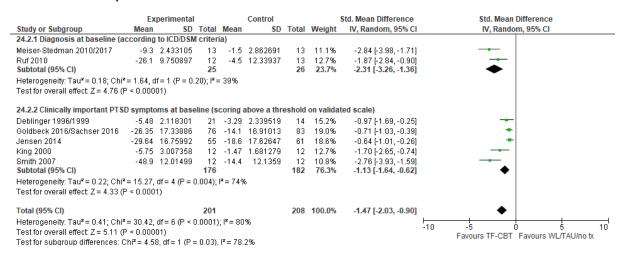
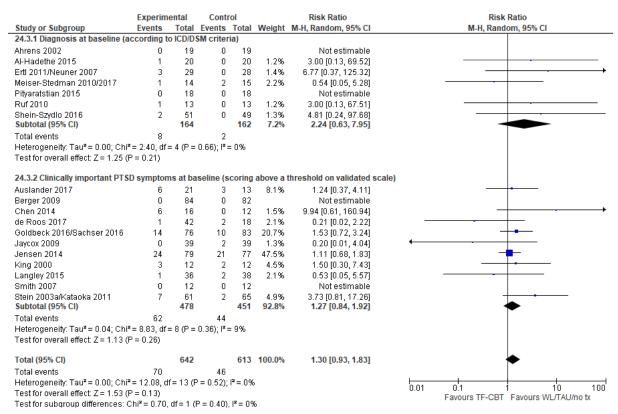


Figure 107: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by trauma type: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 108: Sub-analysis by trauma type: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important

# symptoms/PTSD: PTSD symptomatology self-rated at endpoint (SPTSS/CPSS/CRIES/CRTI/UCLA PTSD-RI/CPTS-RI change score)

| Study or Subgroup  | Ex<br>Mean   | perimental     | Total            | Mean                   | Control  | Total            | Weight        | Std. Mean Difference<br>IV, Random, 95% CI                     | Std. Mean Difference<br>IV, Random, 95% CI           |
|--|--------------|----------------|------------------|------------------------|----------|------------------|---------------|--|--|
| 25.1.1 Motor vehicle collisions  | Weali        | 30             | Total            | Weali                  | 30       | Total            | weight        | IV, Kalluolli, 95% Ci  | IV, Kalluolli, 95% Cl                                |
| Meiser-Stedman 2010/2017   | -24 9        | 6.945142       | 13               | -5.8                   | 10.59481 | 13               | 6.0%          | -2.06 [-3.05, -1.08]   |  |
| Smith 2007   |              | 7.645914       | 12               |                        | 9.626266 | 11               | 4.2%          | -3.65 [-5.06, -2.23]   | <del></del>  |
| Subtotal (95% CI)  |              |                | 25               |                        |          | 24               | 10.3%         | -2.77 [-4.31, -1.23]   | •  |
| Heterogeneity: Tau² = 0.86; Chi²   | = 3.23, 0    | lf = 1 (P = 0. | 07); l² =        | 69%                    |          |                  |               |  |  |
| Test for overall effect: Z = 3.52 (F   | 9 = 0.000    | 4)             |                  |                        |          |                  |               |  |  |
| 25.1.2 Natural disasters   |              |                |                  |                        |          |                  |               |  |  |
| Chen 2014  | -14          | 19.94091       | 10               | 1.55                   | 9.007308 | 12               | 6.4%          | -1.00 [-1.90, -0.10]   |  |
| Pityaratstian 2015   | -1.94        | 9.39987        | 18               | 0.39                   | 9.776293 | 18               | 7.7%          | -0.24 [-0.89, 0.42]  | <u>.</u> †   |
| Subtotal (95% CI)  |              |                | 28               |                        |          | 30               | 14.2%         | -0.55 [-1.29, 0.18]  | •  |
| Heterogeneity: Tau <sup>2</sup> = 0.13; Chi <sup>2</sup><br>Test for overall effect: Z = 1.47 (F |              | If= 1 (P = 0.  | 18); l²=         | : 45%                  |          |                  |               |  |  |
| 25.1.3 Non-sexual violence   |              |                |                  |                        |          |                  |               |  |  |
| Jaycox 2009  |              | 6.890076       | 39               | -1.09                  | 7.633223 | 37               | 8.8%          | -0.36 [-0.81, 0.09]  | <del>-  </del>                                       |
| Stein 2003a/Kataoka 2011   | -15.6        | 5.074692       | 54               | -8                     | 7.007111 | 63               | 9.0%          | -1.22 [-1.62, -0.82]   | _  |
| Subtotal (95% CI)  |              |                | 93               |                        |          | 100              | 17.8%         | -0.80 [-1.64, 0.04]  | •  |
| Heterogeneity: Tau² = 0.32; Chi²<br>Test for overall effect: Z = 1.86 (F                         |              | iτ=1 (P=U.     | UU5); I*         | = 87%                  |          |                  |               |  |  |
| 25.1.4 Mixed   |              |                |                  |                        |          |                  |               |  |  |
| Auslander 2017   | -5.68        | 6.711721       | 15               | 0.8                    | 9.683636 | 10               | 6.8%          | -0.78 [-1.62, 0.05]  | <del></del>  |
| de Roos 2017   |              | 16.22218       | 42               |                        | 15.82135 | 18               | 7.8%          | -1.73 [-2.37, -1.09]   | <del></del>  |
| Goldbeck 2016/Sachser 2016   |              | 9.633047       | 74               |                        | 9.176448 | 82               | 9.3%          | -0.62 [-0.94, -0.30]   | *  |
| Jensen 2014  |              | 6.960165       |                  |                        | 7.633715 | 63               | 9.2%          | -0.74 [-1.11, -0.38]   |  |
| Langley 2015   |              | 9.910537       | 35               |                        | 9.824986 | 36               | 8.5%          | -1.24 [-1.75, -0.73]   | _*   |
| Shein-Szydlo 2016<br>Subtotal (95% CI)   | -23.12       | 8.117503       | 50<br><b>275</b> | -1.94                  | 9.839888 | 49<br><b>258</b> | 8.4%<br>50.1% | -2.40 [-2.92, -1.88]<br>- <b>1.24 [-1.81</b> , - <b>0.68</b> ] | •  |
| Heterogeneity: Tau <sup>2</sup> = 0.42; Chi <sup>2</sup>   | = 40 48      | df = 5 (P < 1  |                  | ): I <sup>2</sup> = 88 | 396      | 250              | 30.170        | -1.24 [-1.01, -0.00]   | •  |
| Test for overall effect: Z = 4.33 (F   |              |                |                  | 7,1 - 00               | ,,,      |                  |               |  |  |
| 25.1.5 Unclear   |              |                |                  |                        |          |                  |               |  |  |
| Al-Hadethe 2015  | -5.05        | 5.638014       | 19               | 2.1                    | 7.251207 | 20               | 7.6%          | -1.07 [-1.75, -0.40]   | -  |
| Subtotal (95% CI)  |              |                | 19               |                        |          | 20               | 7.6%          | -1.07 [-1.75, -0.40]   | •  |
| Heterogeneity: Not applicable  |              |                |                  |                        |          |                  |               |  |  |
| Test for overall effect: Z = 3.11 (F   | 9 = 0.002    | )              |                  |                        |          |                  |               |  |  |
| Total (95% CI)   |              |                | 440              |                        |          | 432              | 100.0%        | -1.21 [-1.59, -0.83]   | •  |
| Heterogeneity: Tau <sup>2</sup> = 0.38; Chi <sup>2</sup>   | = 72.86.     | df = 12 (P <   |                  | )1);                   | 34%      |                  |               |  |  |
| Test for overall effect: Z = 6.18 (F   |              |                |                  |                        |          |                  |               |  | -10 -5 0 5 10<br>Favours TF-CBT Favours WL/TAU/no tx |
| Test for subgroup differences: C   | $hi^2 = 7.3$ | 3, df = 4 (P = | = 0.12),         | $I^2 = 45.4$           | 1%       |                  |               |  | Tarouis III-ODT Tarouis WETAO/III W                  |

Figure 109: Sub-analysis by trauma type: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (CAPS/K-SADS-E: PTSD/ADIS-C:PTSD/CPTSDI; change score)

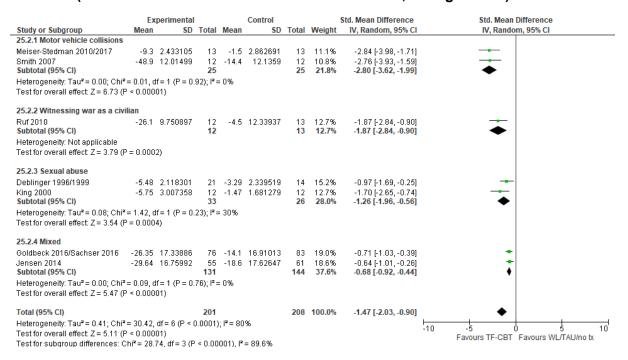
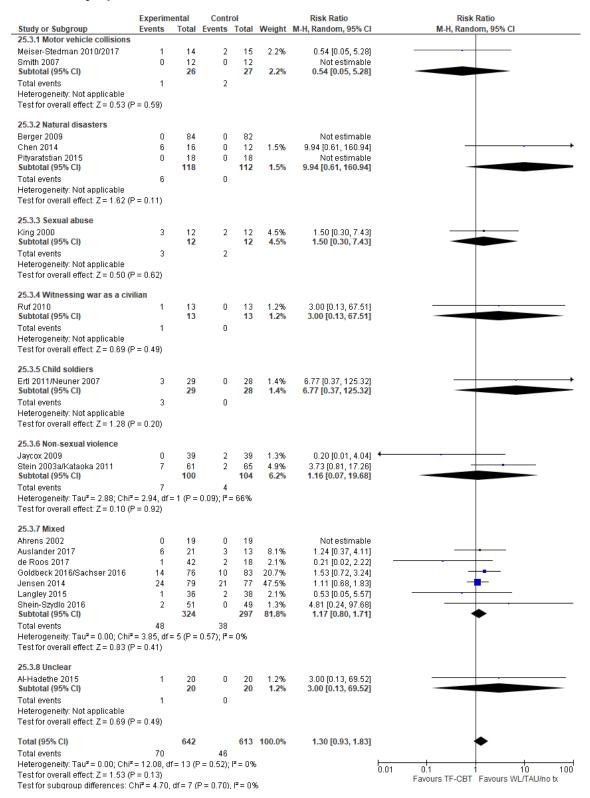


Figure 110: Sub-analysis by trauma type: Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Non-sexual violence



Sub-analysis by specific intervention: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 111: Sub-analysis by specific intervention: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA PTSD-RI/CPSS change score)

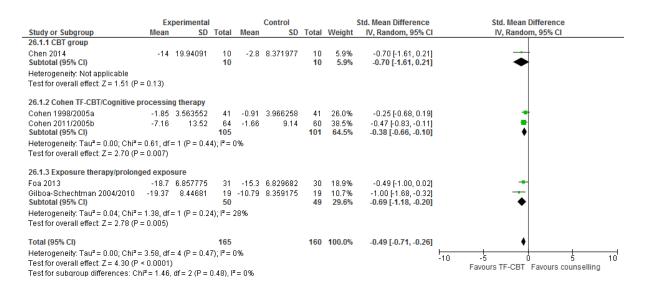
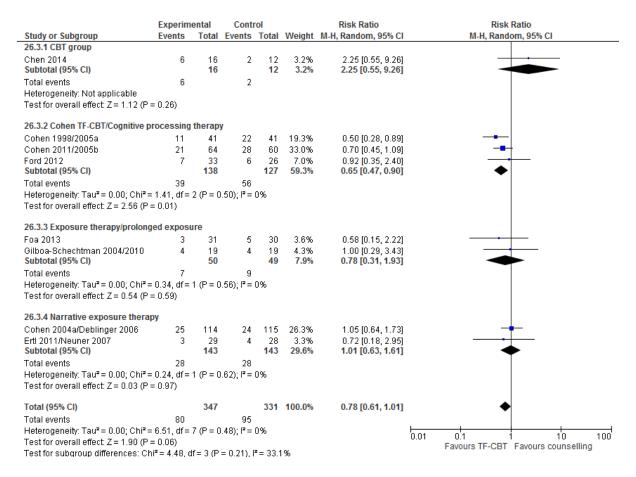


Figure 112: Sub-analysis by specific intervention: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

|   | Ex        | perimental        |                 |         | Control           |                 |                       | Std. Mean Difference  |     | Std. Mean Difference                |    |
|---|-----------|-------------------|-----------------|---------|-------------------|-----------------|-----------------------|---|-----|-------------------------------------|----|
| Study or Subgroup                                 | Mean      | SD                | Total           | Mean    | SD                | Total           | Weight                | IV, Random, 95% CI  |     | IV, Random, 95% CI                  |    |
| 26.2.1 Cohen TF-CBT                               | /Cognitiv | e processi        | ng the          | гару    |                   |                 |                       |   |     |                                     |    |
| Cohen 2011/2005b                                  | -3.31     | 3.48              | 64              | -1.68   | 3.22              | 60              | 43.5%                 | -0.48 [-0.84, -0.13]  |     | -                                   |    |
| Ford 2012<br>Subtotal (95% CI)                    | -24.4     | 13.9318           | 26<br><b>90</b> | -17     | 9.526804          | 20<br><b>80</b> | 26.7%<br><b>70.2%</b> | -0.59 [-1.19, 0.00]<br>- <b>0.51 [-0.82</b> , - <b>0.21</b> ] |     | •                                   |    |
| Heterogeneity: Tau² =<br>Test for overall effect: |           |                   |                 | = 0.75) | ; I² = 0%         |                 |                       |   |     |                                     |    |
| 26.2.2 Exposure ther                              | apy/prol  | onged expo        | sure            |         |                   |                 |                       |   |     |                                     |    |
| Foa 2013  | -20.6     | 6.311747          | 31              | -13.3   | 6.378313          | 30              | 29.8%                 | -1.14 [-1.68, -0.59]  |     | <del></del>                         |    |
| Subtotal (95% CI)                                 |           |                   | 31              |         |                   | 30              | 29.8%                 | -1.14 [-1.68, -0.59]  |     | <b>◆</b>                            |    |
| Heterogeneity: Not ap                             | plicable  |                   |                 |         |                   |                 |                       |   |     |                                     |    |
| Test for overall effect:                          | Z = 4.10  | (P < 0.0001       | )               |         |                   |                 |                       |   |     |                                     |    |
| Total (95% CI)                                    |           |                   | 121             |         |                   | 110             | 100.0%                | -0.71 [-1.10, -0.31]  |     | <b>•</b>                            |    |
| Heterogeneity: Tau² =                             | 0.06; Ch  | $i^2 = 3.94$ , di | f= 2 (P         | = 0.14) | ; I² = 49%        |                 |                       |   | -10 | -5 0 5                              | 10 |
| Test for overall effect:                          | Z = 3.49  | (P = 0.0006)      | 5)              |         |                   |                 |                       |   | -10 | Favours TF-CBT Favours counsellin   |    |
| Test for subaroup diff                            | erences:  | $Chi^2 = 3.84$    | l. df = 1       | (P = 0. | 05), $I^2 = 73.9$ | 3%              |                       |   |     | Tavouis III-ODT Tavouis coulisellii | 9  |

Figure 113: Sub-analysis by specific intervention: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by format: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 114: Sub-analysis by format: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important

#### symptoms/PTSD: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA PTSD-RI/CPSS change score)

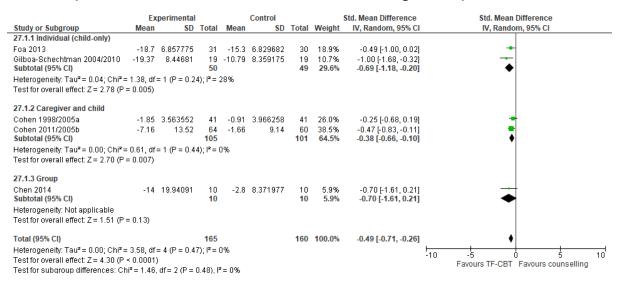
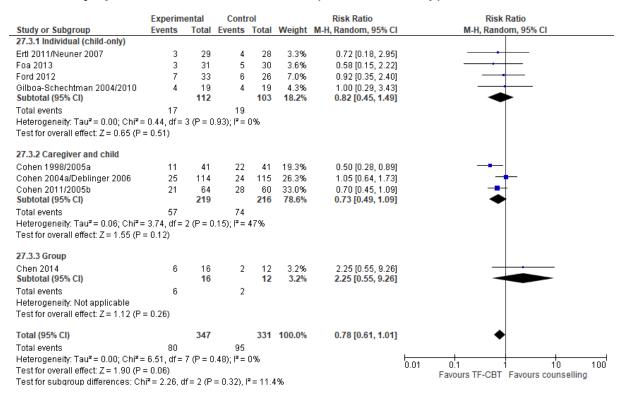


Figure 115: Sub-analysis by format: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

|  | Ex       | perimental |                 |         | Control   |                 |                       | Std. Mean Difference                                  |     | Std. Mean Difference                 |                   |
|--|----------|------------|-----------------|---------|-----------|-----------------|-----------------------|---|-----|--------------------------------------|-------------------|
| Study or Subgroup  | Mean     | SD         | Total           | Mean    | SD        | Total           | Weight                | IV, Random, 95% CI                                    |     | IV, Random, 95% CI                   |                   |
| 27.2.1 Individual (chi   | ld-only) |            |                 |         |           |                 |                       |   |     |                                      |                   |
| Foa 2013   | -20.6    | 6.311747   | 31              | -13.3   | 6.378313  | 30              | 29.8%                 | -1.14 [-1.68, -0.59]                                  |     |                                      |                   |
| Ford 2012<br>Subtotal (95% CI)   | -24.4    | 13.9318    | 26<br><b>57</b> | -17     | 9.526804  | 20<br><b>50</b> | 26.7%<br><b>56.5%</b> | -0.59 [-1.19, 0.00]<br>- <b>0.88 [-1.41, -0.35]</b>   |     | •                                    |                   |
| Heterogeneity: Tau² =<br>Test for overall effect:                                      |          |            | •               | = 0.19) | ; I²= 42% |                 |                       |   |     |                                      |                   |
| 27.2.2 Caregiver and   | child    |            |                 |         |           |                 |                       |   |     |                                      |                   |
| Cohen 2011/2005b<br>Subtotal (95% CI)  | -3.31    | 3.48       | 64<br><b>64</b> | -1.68   | 3.22      | 60<br><b>60</b> | 43.5%<br><b>43.5%</b> | -0.48 [-0.84, -0.13]<br>- <b>0.48 [-0.84, -0.13</b> ] |     | •                                    |                   |
| Heterogeneity: Not ap<br>Test for overall effect:                                      | •        |            |                 |         |           |                 |                       |   |     |                                      |                   |
| Total (95% CI)   |          |            | 121             |         |           | 110             | 100.0%                | -0.71 [-1.10, -0.31]                                  |     | •                                    |                   |
| Heterogeneity: Tau <sup>2</sup> =<br>Test for overall effect:<br>Test for subgroup dif | Z = 3.49 | (P = 0.000 | 5)              | ·       |           | 7%              |                       |   | -10 | -5 0 :<br>Favours TF-CBT Favours cou | 5 10<br>Inselling |

Figure 116: Sub-analysis by format: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by age range: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 117: Sub-analysis by age range: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA PTSD-RI/CPSS change score)

|  | Ex                     | perimental          |   |                 | Control                          |                             |  | Std. Mean Difference   | Std. Mean Difference                             |
|--|------------------------|---------------------|---|-----------------|----------------------------------|-----------------------------|--|--|--|
| Study or Subgroup  | Mean                   | SD                  | Total                                   | Mean            | SD                               | Total                       | Weight                                 | IV, Random, 95% CI   | IV, Random, 95% CI                               |
| 28.1.1 Age range includes age 7  | and und                | ег                  |   |                 |                                  |                             |  |  |  |
| Cohen 1998/2005a   | -1.85                  | 3.563552            | 41                                      | -0.91           | 3.966258                         | 41                          | 26.0%                                  | -0.25 [-0.68, 0.19]  | <del>-</del>                                     |
| Cohen 2011/2005b   | -7.16                  | 13.52               | 64                                      | -1.66           | 9.14                             | 60                          | 38.5%                                  | -0.47 [-0.83, -0.11]   | <u>₹</u>   |
| Subtotal (95% CI)  |                        |                     | 105                                     |                 |                                  | 101                         | 64.5%                                  | -0.38 [-0.66, -0.10]   | <b>♦</b>   |
| Heterogeneity: Tau² = 0.00; Chi² =   | 0.61, df               | = 1 (P = 0.4        | 4); $  ^2 = 1$                          | 0%              |                                  |                             |  |  |  |
| Test for overall effect: Z = 2.70 (P =   | = 0.007)               |                     |   |                 |                                  |                             |  |  |  |
| 28.1.2 Age range only includes of<br>Chen 2014<br>Foa 2013<br>Gilboa-Schechtman 2004/2010<br>Subtotal (95% CI)<br>Heterogeneity: Tau² = 0.00; Chi² = | -14<br>-18.7<br>-19.37 | 6.857775<br>8.44681 | 10<br>31<br>19<br><b>60</b><br>0);  ² = | -15.3<br>-10.79 | 8.371977<br>6.829682<br>8.359175 | 10<br>30<br>19<br><b>59</b> | 5.9%<br>18.9%<br>10.7%<br><b>35.5%</b> | -0.70 [-1.61, 0.21]<br>-0.49 [-1.00, 0.02]<br>-1.00 [-1.68, -0.32]<br>-0.68 [-1.05, -0.31] |  |
| Test for overall effect: Z = 3.58 (P =   | = 0.0003               | )                   |   |                 |                                  |                             |  |  |  |
| Total (95% CI)  Heterogeneity: Tau* = 0.00; Chi* =  Test for overall effect: Z = 4.30 (P of Test for subgroup differences: Ch                        | < 0.0001               | )                   |   |                 | 6                                | 160                         | 100.0%                                 | -0.49 [-0.71, -0.26]   | -10 -5 0 5 10 Favours TF-CBT Favours counselling |

Figure 118: Sub-analysis by age range: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important

## symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

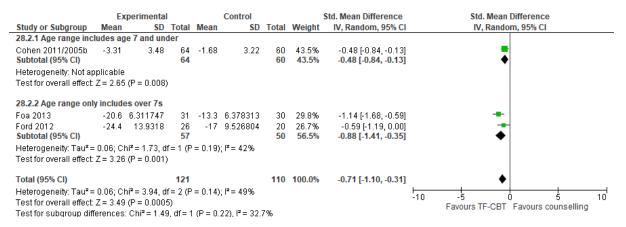
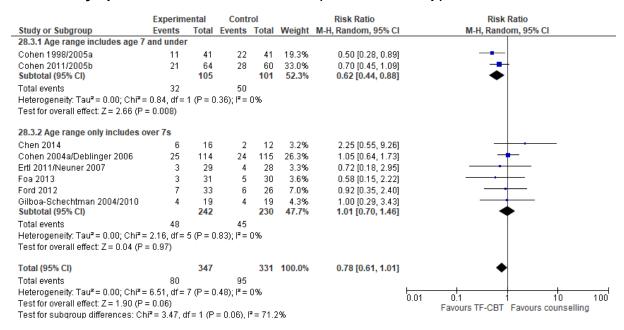


Figure 119: Sub-analysis by age range: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 120: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of

## clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA PTSD-RI/CPSS change score)

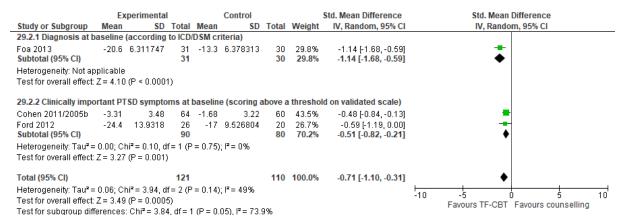
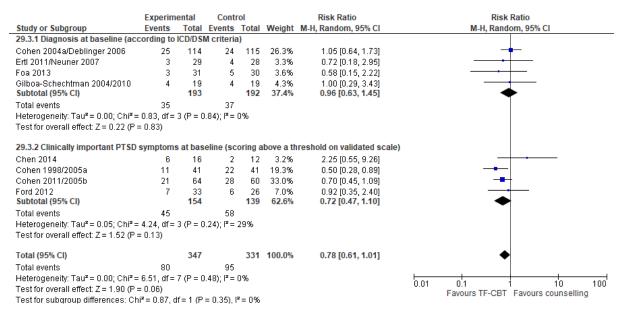


Figure 121: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

|                             | Ex                         | cperimenta       | ı               |         | Control           |                 |                       | Std. Mean Difference   |     | Std. Mean Differ    | ence                 |    |
|-----------------------------|----------------------------|------------------|-----------------|---------|-------------------|-----------------|-----------------------|--|-----|---------------------|----------------------|----|
| Study or Subgro             | oup Mean                   | SD               | Total           | Mean    | SD                | Total           | Weight                | IV, Random, 95% CI   |     | IV, Random, 95      | % CI                 |    |
| 29.2.1 Diagnosis            | s at baseline              | (according       | to ICD          | DSM ci  | riteria)          |                 |                       |  |     |                     |                      |    |
| Foa 2013<br>Subtotal (95% C |                            | 6.311747         | 31<br><b>31</b> | -13.3   | 6.378313          | 30<br><b>30</b> | 29.8%<br><b>29.8%</b> | -1.14 [-1.68, -0.59]<br>- <b>1.14</b> [- <b>1.68</b> , - <b>0.59</b> ] |     | •                   |                      |    |
| Heterogeneity: N            | Not applicable             | !                |                 |         |                   |                 |                       |  |     |                     |                      |    |
| Test for overall e          | effect: Z = 4.10           | ) (P < 0.000     | 1)              |         |                   |                 |                       |  |     |                     |                      |    |
| 29.2.2 Clinically           | important P1               | SD sympto        | oms at          | baselin | e (scoring a      | above a         | threshole             | d on validated scale)  |     |                     |                      |    |
| Cohen 2011/200              | 05b -3.31                  | 3.48             | 64              | -1.68   | 3.22              | 60              | 43.5%                 | -0.48 [-0.84, -0.13]   |     | -                   |                      |    |
| Ford 2012                   | -24.4                      | 13.9318          | 26              | -17     | 9.526804          | 20              | 26.7%                 | -0.59 [-1.19, 0.00]  |     | <del></del>         |                      |    |
| Subtotal (95% C             | (I)                        |                  | 90              |         |                   | 80              | 70.2%                 | -0.51 [-0.82, -0.21]   |     | <b>♦</b>            |                      |    |
| Heterogeneity: T            | Γau² = 0.00; C             | $hi^2 = 0.10, c$ | if = 1 (P       | = 0.75) | ; I² = 0%         |                 |                       |  |     |                     |                      |    |
| Test for overall e          | effect: Z = 3.27           | ' (P = 0.001     | )               |         |                   |                 |                       |  |     |                     |                      |    |
| Total (95% CI)              |                            |                  | 121             |         |                   | 110             | 100.0%                | -0.71 [-1.10, -0.31]   |     | •                   |                      |    |
| Heterogeneity: T            | Tau <sup>2</sup> = 0.06; C | $hi^2 = 3.94, c$ | lf = 2 (P       | = 0.14) | ; I² = 49%        |                 |                       |  | 4.0 | -5 0                | <del>_</del>         | 11 |
| Test for overall e          | effect: Z = 3.49           | P = 0.000        | 5)              |         |                   |                 |                       |  | -10 | Favours TF-CBT Favo | O<br>ure councelling |    |
| Test for subarou            | up differences             | : Chi² = 3.8     | 4. df = 1       | (P = 0. | 05), $I^2 = 73.9$ | 9%              |                       |  |     | ravouis ir-CB1 ravo | urs couriseiling     | 1  |

Figure 122: Sub-analysis by diagnostic status at baseline: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



Sub-analysis by trauma type: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 123: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA P Sub-analysis by trauma type: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: TSD-RI/CPSS change score)

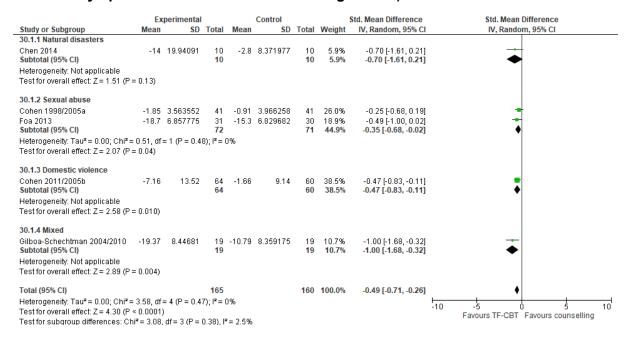


Figure 124: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA P Sub-analysis by trauma type: Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated at endpoint (K-SADS-PL: PTSD/CPSS-I/CAPS; change score)

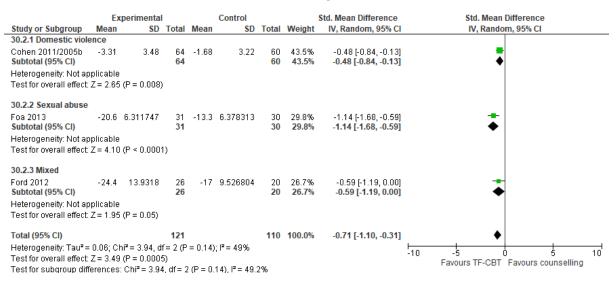
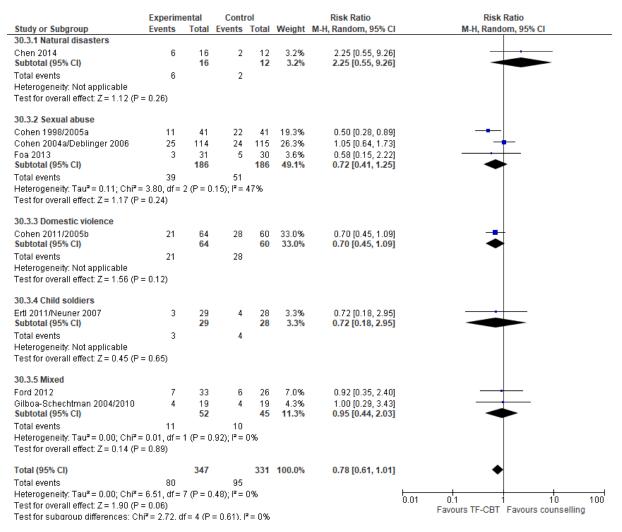


Figure 125: PTSD symptomatology self-rated at endpoint (CRIES/TSCC-PTSD/UCLA P Sub-analysis by trauma type: Trauma-focused CBT versus supportive

## counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



#### Psychological: Non-trauma-focused CBT

Non-trauma focused CBT (+ TAU) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 126: Non-trauma focused CBT (+ TAU) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms

# (Adolescent Psychopathology Scale: Axis I - Major Depression; change score); Multiple incident index trauma

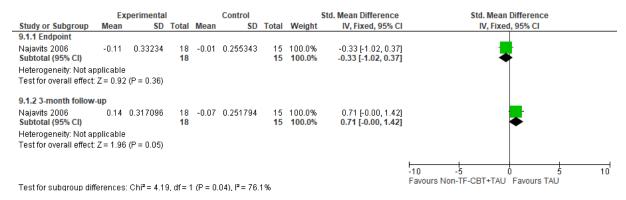


Figure 127: Non-trauma focused CBT (+ TAU) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Substance use disorder symptoms (Adolescent Psychopathology Scale: Axis I - Substance Use Disorder; change score); Multiple incident index trauma

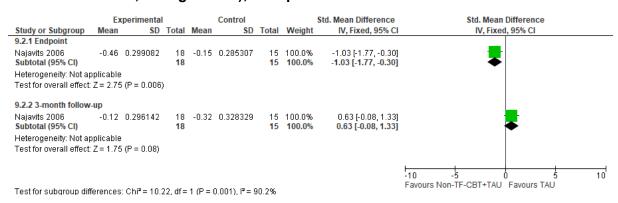
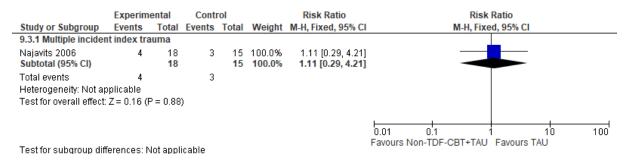


Figure 128: Non-trauma focused CBT (+ TAU) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



#### Psychological: Psychodynamic therapies

Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 129: Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (DC 0-3; change score)

|   | Ex    | perimental | ı               |      | Control  |                 |        | Std. Mean Difference                                  | Std. Mean   | Difference |    |
|---|-------|------------|-----------------|------|----------|-----------------|--------|---|-------------|------------|----|
| Study or Subgroup   | Mean  | SD         | Total           | Mean | SD       | Total           | Weight | IV, Fixed, 95% CI                                     | IV, Fixed   | I, 95% CI  |    |
| 11.1.1 Multiple incident index trauma   |       |            |                 |      |          |                 |        |   |             |            |    |
| Lieberman 2005/2006/Ghosh Ippen 2011<br>Subtotal (95% CI)                       | -3.61 | 2.326929   | 36<br><b>36</b> | -0.4 | 3.028795 | 29<br><b>29</b> |        | -1.19 [-1.72, -0.66]<br>- <b>1.19 [-1.72, -0.66</b> ] |             |            |    |
| Heterogeneity: Not applicable<br>Test for overall effect: Z = 4.38 (P < 0.0001) |       |            |                 |      |          |                 |        |   |             |            |    |
|   |       |            |                 |      |          |                 |        |   | -10 -5 I    | ) 5        | 10 |
| Test for subgroup differences: Not applicabl                                    | е     |            |                 |      |          |                 |        |   | Favours CFF | Favours FT |    |

Figure 130: Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems (CBCL total; change score); Multiple incident index trauma

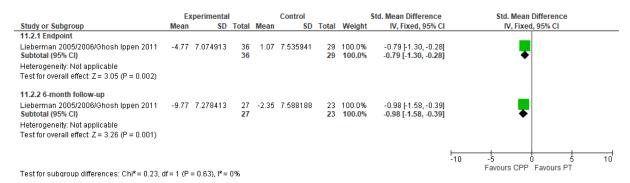


Figure 131: Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Discontinuation (loss to follow-up)

|   | Experim  | ental           | Cont   | rol             |                         | Risk Ratio                             | Risk Ratio                                |
|---|----------|-----------------|--------|-----------------|-------------------------|--|---|
| Study or Subgroup   | Events   | Total           | Events | Total           | Weight                  | M-H, Fixed, 95% CI                     | M-H, Fixed, 95% CI                        |
| 11.3.1 Multiple incident index trauma   |          |                 |        |                 |                         |  |   |
| Lieberman 2005/2006/Ghosh Ippen 2011<br>Subtotal (95% CI)                               | 6        | 42<br><b>42</b> | 4      | 33<br><b>33</b> | 100.0%<br><b>100.0%</b> | 1.18 [0.36, 3.84]<br>1.18 [0.36, 3.84] | <b>*</b>                                  |
| Total events Heterogeneity: Not applicable Test for overall effect: Z = 0.27 (P = 0.78) | 6        |                 | 4      |                 |                         |  |   |
| Test for subgroup differences: Not applicable   | <b>.</b> |                 |        |                 |                         |  | 0.01 0.1 10 100<br>Favours CPP Favours PT |

#### Psychological: Eye movement desensitization and reprocessing (EMDR)

## EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 132: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at endpoint (CRTI/CRIES change score)

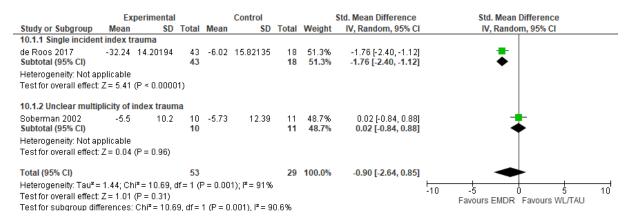


Figure 133: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated at 2-month follow-up (CRIES change score)

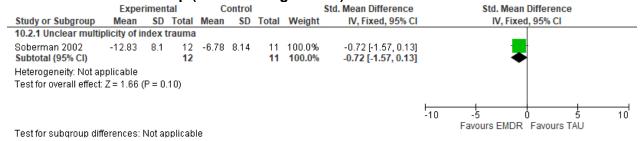


Figure 134: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (PTSS-C change score)

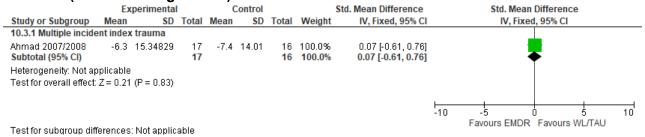


Figure 135: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems (SDQ-A change score)

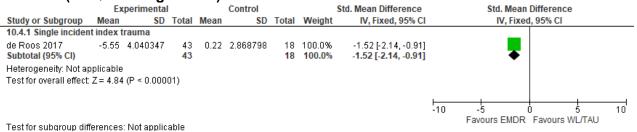


Figure 136: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Quality of life (KIDSCREEN-27: Global HRQoL T-scores; change score)

|   | Ex         | cperimental   | ı               |      | Control  |                 |                          | Std. Mean Difference                          | Std. Mean Difference        |
|---|------------|---------------|-----------------|------|----------|-----------------|--------------------------|---|-----------------------------|
| Study or Subgroup                               | Mean       | SD            | Total           | Mean | SD       | Total           | Weight                   | IV, Fixed, 95% CI                             | IV, Fixed, 95% CI           |
| 10.5.1 Single incider                           | nt index t | trauma        |                 |      |          |                 |                          |   |                             |
| de Roos 2017<br>Subtotal (95% CI)               | 10.23      | 11.11799      | 43<br><b>43</b> | 1.07 | 11.14915 | 18<br><b>18</b> | 100.0%<br><b>100.0</b> % | 0.81 [0.24, 1.38]<br><b>0.81 [0.24, 1.38]</b> | <b>.</b>                    |
| Heterogeneity: Not a<br>Test for overall effect |            |               | )               |      |          |                 |                          |   |                             |
|   |            |               |                 |      |          |                 |                          |   | -10 -5 0 5 10               |
| Test for subgroup dit                           | fferences  | s: Not applic | able            |      |          |                 |                          |   | Favours WL/TAU Favours EMDR |

Figure 137: EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)

|                                      | Experim          | ental                 | Conti         | rol             |                         | Risk Ratio                                    | Risk Ratio                  |     |
|--------------------------------------|------------------|-----------------------|---------------|-----------------|-------------------------|---|-----------------------------|-----|
| Study or Subgroup                    | Events           | Total                 | <b>Events</b> | Total           | Weight                  | M-H, Random, 95% CI                           | M-H, Random, 95% CI         |     |
| 10.6.1 Single incide                 | nt index tra     | uma                   |               |                 |                         |   |                             |     |
| de Roos 2017<br>Subtotal (95% CI)    | 1                | 43<br><b>43</b>       | 2             | 18<br><b>18</b> | 30.5%<br><b>30.5%</b>   | 0.21 [0.02, 2.17]<br><b>0.21 [0.02, 2.17]</b> |                             |     |
| Total events                         | 1                |                       | 2             |                 |                         |   |                             |     |
| Heterogeneity: Not a                 | pplicable        |                       |               |                 |                         |   |                             |     |
| Test for overall effect              | t: Z = 1.31 (F   | P = 0.19              | )             |                 |                         |   |                             |     |
| 10.6.2 Multiple incid                | lent index t     | rauma                 |               |                 |                         |   |                             |     |
| Ahmad 2007/2008<br>Subtotal (95% CI) | 0                | 17<br><b>17</b>       | 0             | 16<br><b>16</b> |                         | Not estimable<br><b>Not estimable</b>         | l l                         |     |
| Total events                         | 0                |                       | 0             |                 |                         |   |                             |     |
| Heterogeneity: Not a                 | pplicable        |                       |               |                 |                         |   |                             |     |
| Test for overall effect              | t: Not applic    | able                  |               |                 |                         |   |                             |     |
| 10.6.3 Unclear multi                 | iplicity of in   | dex tra               | uma           |                 |                         |   |                             |     |
| Soberman 2002                        | 4                | 14                    | 4             | 15              | 69.5%                   | 1.07 [0.33, 3.48]                             | <del></del>                 |     |
| Subtotal (95% CI)                    |                  | 14                    |               | 15              | 69.5%                   | 1.07 [0.33, 3.48]                             | -                           |     |
| Total events                         | 4                |                       | 4             |                 |                         |   |                             |     |
| Heterogeneity: Not a                 | pplicable        |                       |               |                 |                         |   |                             |     |
| Test for overall effect              | t: $Z = 0.11$ (F | P = 0.91              | )             |                 |                         |   |                             |     |
| Total (95% CI)                       |                  | 74                    |               | 49              | 100.0%                  | 0.65 [0.15, 2.88]                             |                             |     |
| Total events                         | 5                |                       | 6             |                 |                         |   |                             |     |
| Heterogeneity: Tau <sup>2</sup> :    | = 0.46; Chi²     | ²= 1.52,              | df = 1 (P     | = 0.22          | ); I <sup>z</sup> = 34% | ,<br>,  | 0.01 0.1 1 10               | 100 |
| Test for overall effect              | t: Z = 0.57 (F   | P = 0.57              | )             |                 |                         |   | Favours EMDR Favours WL/TA  |     |
| Test for subgroup di                 | fferences: 0     | Chi <sup>z</sup> = 1. | 50, df = 1    | (P = 0.         | .22), $I^2 = 3$         | 3.1%  | TAVOUIS EMDIT TAVOUIS WETTA |     |

### Psychological: Combined somatic and cognitive therapies

Combined somatic and cognitive therapies versus no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 138: Combined somatic and cognitive therapies versus no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated (SPTSS change score); Unclear multiplicity of index trauma

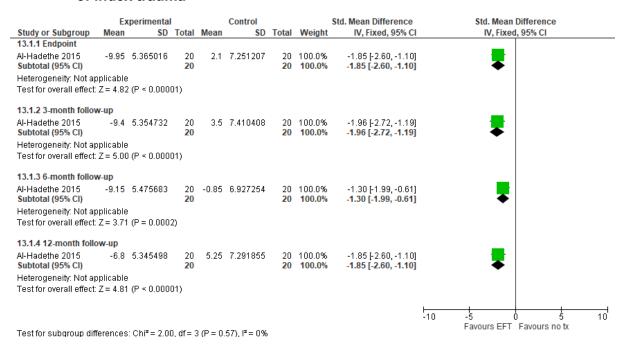


Figure 139: Combined somatic and cognitive therapies versus no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

## Anxiety symptoms (HADS-A change score); Unclear multiplicity of index trauma

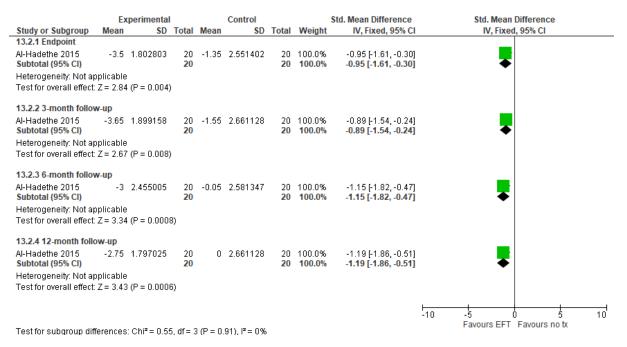
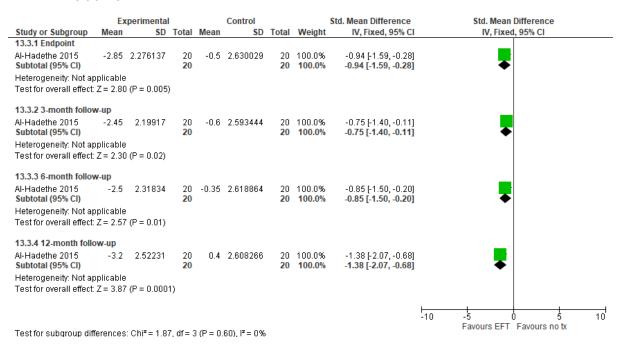


Figure 140: Combined somatic and cognitive therapies versus no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Depression symptoms (HADS-D change score); Unclear multiplicity of index trauma



#### **Psychological: Supportive counselling**

Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 141: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated (CRIES change score); Single incident index trauma

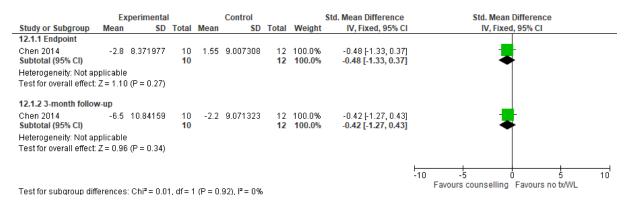


Figure 142: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (CAPS change score); Multiple incident index trauma

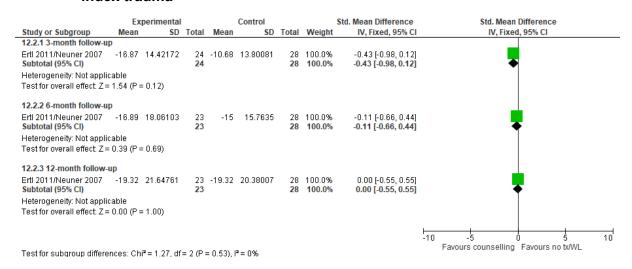


Figure 143: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Remission at

# 12-month follow-up (number of people no longer meeting diagnostic criteria for PTSD)

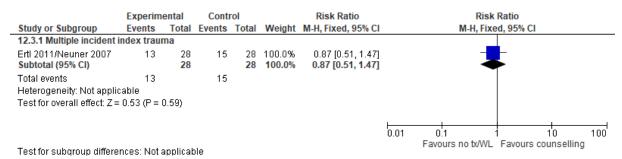


Figure 144: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at endpoint (CES-D change score)

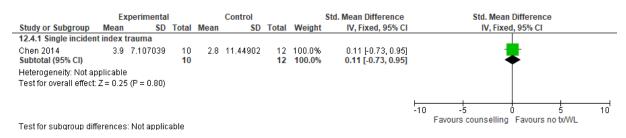


Figure 145: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 3-month follow-up (CES-D/MINI:Depression change score)

|   | E        | cperimental | ı               |             | Control    |                 |                       | Std. Mean Difference                                 |     | Std. Mean Difference                 |
|---|----------|-------------|-----------------|-------------|------------|-----------------|-----------------------|--|-----|--------------------------------------|
| Study or Subgroup   | Mean     | SD          | Total           | Mean        | SD         | Total           | Weight                | IV, Random, 95% CI                                   |     | IV, Random, 95% CI                   |
| 12.5.1 Single incident in                                 | dex trau | ıma         |                 |             |            |                 |                       |  |     |                                      |
| Chen 2014<br>Subtotal (95% CI)                            | -4.8     | 6.750741    | 10<br><b>10</b> | -0.4        | 11.33446   | 12<br><b>12</b> | 30.8%<br><b>30.8%</b> | -0.44 [-1.29, 0.41]<br>- <b>0.44 [-1.29, 0.41]</b>   |     | <del>-</del>                         |
| Heterogeneity: Not applic                                 |          | - 0.24)     |                 |             |            |                 |                       |  |     |                                      |
| Test for overall effect: Z=                               | 1.02 (P  | = 0.31)     |                 |             |            |                 |                       |  |     |                                      |
| 12.5.2 Multiple incident i                                | ndex tra | auma        |                 |             |            |                 |                       |  |     |                                      |
| Ertl 2011/Neuner 2007<br>Subtotal (95% CI)                | 0.39     | 1.957958    | 24<br><b>24</b> | 1.97        | 1.898894   | 28<br><b>28</b> | 69.2%<br>69.2%        | -0.81 [-1.38, -0.24]<br>- <b>0.81 [-1.38, -0.24]</b> |     | •                                    |
| Heterogeneity: Not applic                                 |          |             |                 |             |            |                 |                       | ,,   |     | Ť                                    |
| Test for overall effect: Z=                               | 2.78 (P  | = 0.005)    |                 |             |            |                 |                       |  |     |                                      |
| Total (95% CI)  |          |             | 34              |             |            | 40              | 100.0%                | -0.70 [-1.17, -0.22]                                 |     | •                                    |
| Heterogeneity: Tau <sup>2</sup> = 0.0                     | •        |             | 1 (P = 0        | 0.49); l² : | = 0%       |                 |                       |  | -10 | -5 0 5 10                            |
| Test for overall effect: Z =<br>Test for subgroup differe |          |             | f = 1 /D        | - 0.40\     | IZ — 0.0%  |                 |                       |  |     | Favours counselling Favours no tx/WL |
| restroi sundroup umere                                    | nces. O  | m = 0.45, u | i – i (r        | - 0.43)     | , 1 - 0 70 |                 |                       |  |     |                                      |

Figure 146: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 6-month follow-up (MINI:Depression change score)

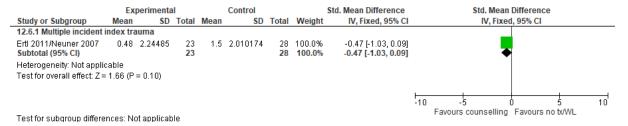


Figure 147: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms at 12-month follow-up (MINI:Depression change score)

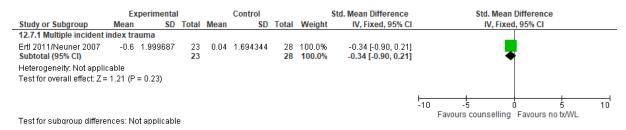


Figure 148: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Functional impairment (CAPS: Functional impairment; change score); Multiple incident index trauma

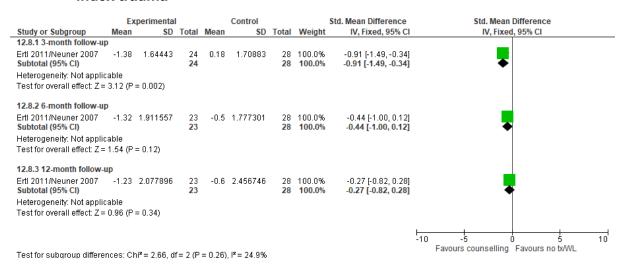
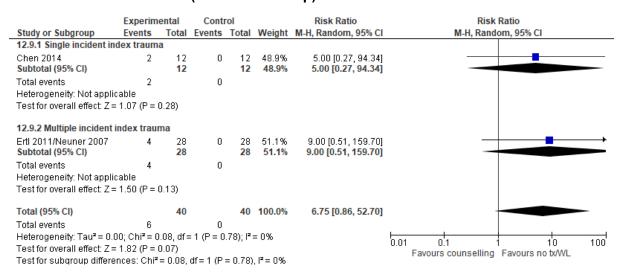


Figure 149: Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD:

Discontinuation (loss to follow-up)



### Psychological: Parent training/family intervention

Parent training (CBT with parent-only) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 150: Parent training (CBT with parent-only) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

## symptomatology clinician-rated (K-SADS-E: PTSD; change score); Multiple incident index trauma

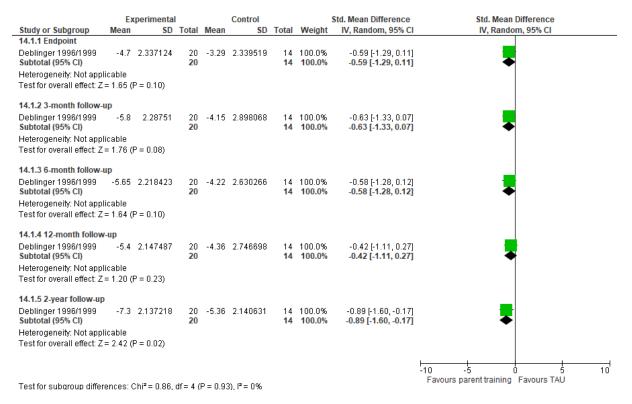
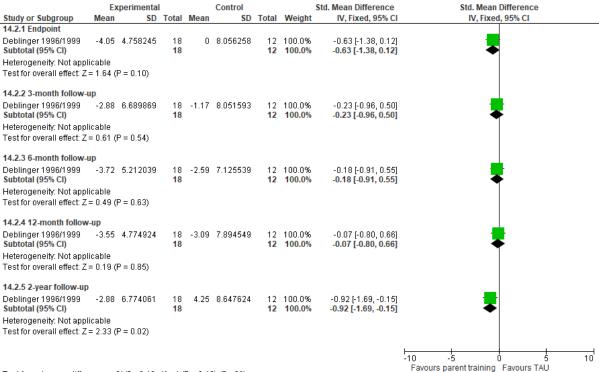


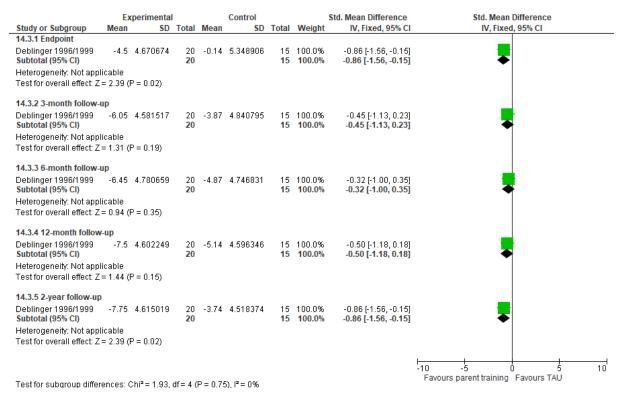
Figure 151: Parent training (CBT with parent-only) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional

# and behavioural problems-Externalizing (CBCL: Externalizing; change score); Multiple incident index trauma



Test for subgroup differences:  $Chi^2 = 3.40$ , df = 4 (P = 0.49),  $I^2 = 0\%$ 

Figure 152: Parent training (CBT with parent-only) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms (CDI change score); Multiple incident index trauma



Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 153: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology clinician-rated (ADIS-C: PTSD; change score); Multiple incident index trauma

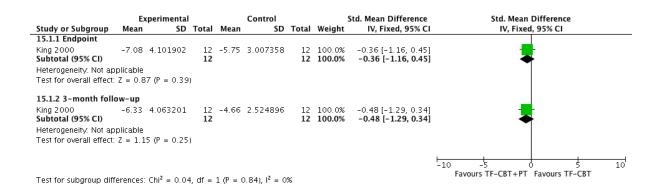


Figure 154: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms (RCMAS; change score); Multiple incident index trauma

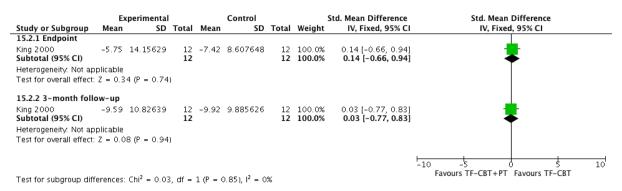


Figure 155: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Depression symptoms (CDI; change score); Multiple incident index trauma

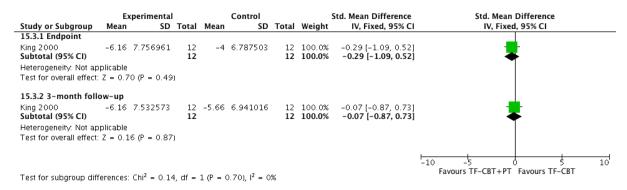


Figure 156: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Internalizing (CBCL: Internalizing; change score); Multiple incident index trauma

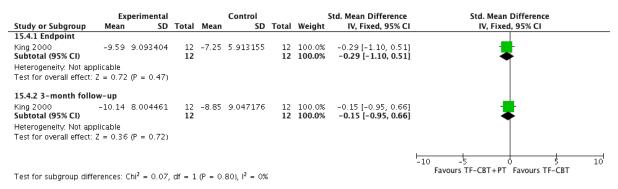


Figure 157: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Emotional and behavioural problems-Externalizing (CBCL: Externalizing; change score); Multiple incident index trauma

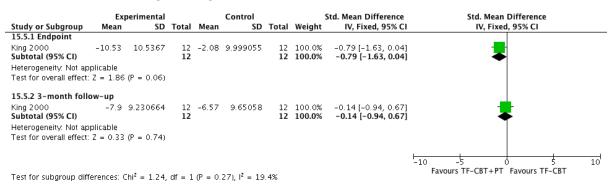


Figure 158: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Global functioning (GAF; change score); Multiple incident index trauma

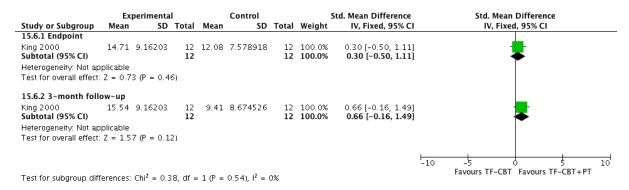
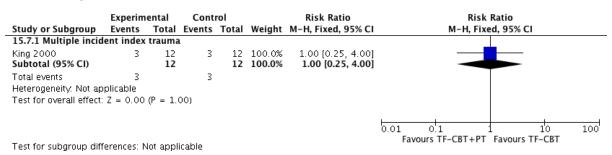


Figure 159: Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



### Family therapy versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 160: Family therapy versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology (UCLA PTSD-RI; change score)

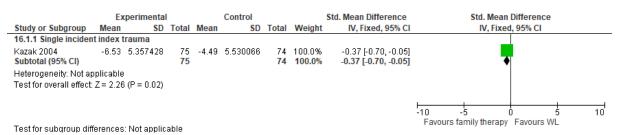


Figure 161: Family therapy versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Anxiety symptoms (RCMAS; T-scores change score)

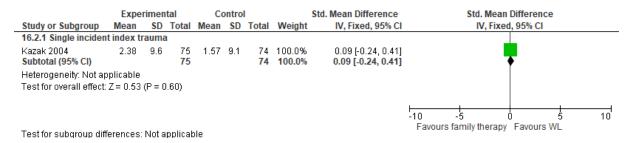
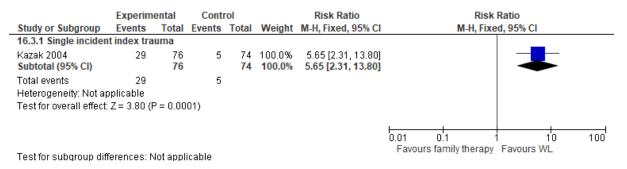


Figure 162: Family therapy versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)



#### Psychological: Play therapy

Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling

Figure 163: Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling: PTSD symptomatology self-rated (CRIES change score)

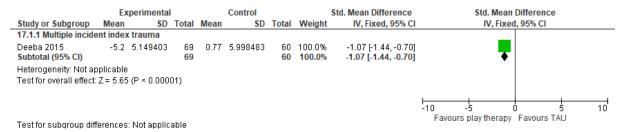


Figure 164: Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling: Anxiety symptoms (SCASp; change score)

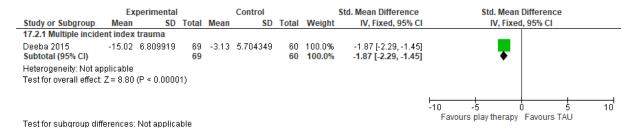


Figure 165: Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling: Depression symptoms (SMFQp; change score)

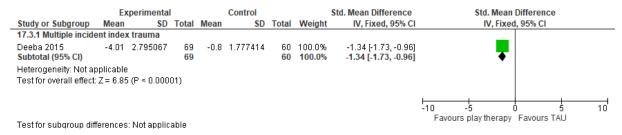
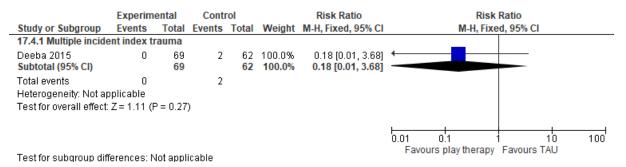


Figure 166: Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling:

Discontinuation (loss to follow-up)



Play therapy versus trauma-focused CBT for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 167: Play therapy versus trauma-focused CBT for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated (UCLA PTSD-RI; change score)

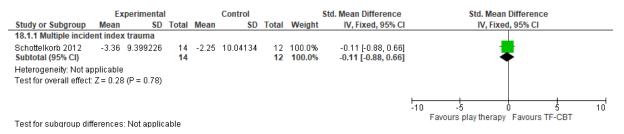
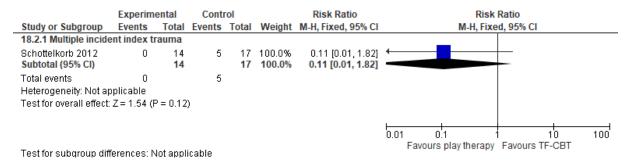


Figure 168: Play therapy versus trauma-focused CBT for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)

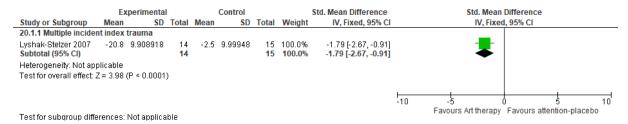


Psychosocial: Art therapy

Art therapy (+ TAU) versus attention-placebo (+ TAU) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 169: Art therapy (+ TAU) versus attention-placebo (+ TAU) for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD

# symptomatology clinician-rated (UCLA PTSD-RI adminstered via structured interview format; change score)



#### **Psychosocial: Meditation**

### Meditation versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

Figure 170: Meditation versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: PTSD symptomatology self-rated (HTQ change score)

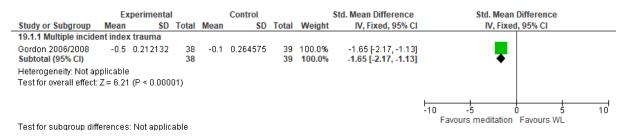
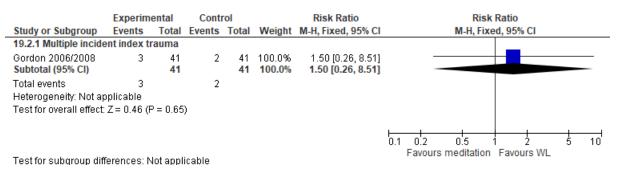


Figure 171: Meditation versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD: Discontinuation (loss to follow-up)





### **Appendix F- GRADE tables**

GRADE tables for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

**Psychological: Trauma-focused CBT** 

Trauma-focused CBT versus meditation for the early treatment (1-3 months) of clinically important symptoms/PTSD

|               |                      |                                   |                             |                            |                              | •                     |                      |                 |                             |   |                 | l and the second se |
|---------------|----------------------|-----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------|----------------------|-----------------|-----------------------------|---|-----------------|--|
| 0             |                      |                                   |                             |                            |                              |                       | No. of con-          |                 | F654                        |   |                 |  |
| Quality       | assessment           |                                   |                             |                            |                              |                       | No of pat            |                 | Effect                      |   |                 |  |
| No of studies | Design               | Risk<br>of<br>bias                | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s | Trauma - focused CBT | Meditatio<br>n  | Relativ<br>e<br>(95%<br>CI) | Absolute  | Quality         | Importanc<br>e   |
| PTSD sy       | /mptomatolog         | gy clinicia                       | n-rated at 1-mor            | ith follow-up (fo          | ollow-up meai                | n 1 months; meas      | ured with:           | CPTS-RI ch      | ange score                  | e; Better ind   | icated by lower | values)  |
| 1             | randomised<br>trials | no<br>seriou<br>s risk<br>of bias | no serious<br>inconsistency | no serious indirectness    | very<br>serious <sup>1</sup> | none                  | 16                   | 15              | -                           | SMD 0.15<br>lower<br>(0.85<br>lower to<br>0.56<br>higher) | LOW             | CRITICAL   |
| PTSD sy       | /mptomatolog         | gy clinicia                       | n-rated at 6-mor            | ith follow-up (fo          | ollow-up meai                | n 6 months; meas      | ured with:           | CPTS-RI ch      | ange score                  | e; Better ind   | icated by lower | values)  |
| 1             | randomised<br>trials | no<br>seriou<br>s risk<br>of bias | no serious inconsistency    | no serious<br>indirectness | very<br>serious <sup>1</sup> | none                  | 16                   | 14              | -                           | SMD 0.12<br>higher<br>(0.6 lower<br>to 0.83<br>higher)    | LOW             | CRITICAL   |
| Diagnos       | is at 1-month        | follow-up                         | o (follow-up mea            | n 1 months; as             | sessed with:                 | Number of people      | who met o            | criteria for a  | diagnosis                   | of PTSD)  |                 |  |
| 1             | randomised<br>trials | no<br>seriou                      | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>1</sup> | none                  | 4/16<br>(25%)        | 5/15<br>(33.3%) | RR<br>0.75                  | 83 fewer<br>per 1000<br>(from 250                         | LOW             | CRITICAL   |

| Quality a     | assessment           |                                   |                             |                         |                              |                       | No of pat            | ients           | Effect                          |   |          |                |
|---------------|----------------------|-----------------------------------|-----------------------------|-------------------------|------------------------------|-----------------------|----------------------|-----------------|---------------------------------|---|----------|----------------|
| No of studies | Design               | Risk<br>of<br>bias                | Inconsistenc<br>y           | Indirectnes<br>s        | Imprecisio<br>n              | Other consideration s | Trauma - focused CBT | Meditatio<br>n  | Relativ<br>e<br>(95%<br>CI)     | Absolute  | Quality  | Importanc<br>e |
|               |                      | s risk<br>of bias                 |                             |                         |                              |                       |                      |                 | (0.25 to<br>2.28)               | fewer to<br>427<br>more)                                      |          |                |
| Diagnos       | is at 6-month        | follow-up                         | o (follow-up mea            | n 6 months; as          | sessed with:                 | Number of people      | who met o            | criteria for a  | diagnosis                       | of PTSD)  |          |                |
| 1             | randomised<br>trials | no<br>seriou<br>s risk<br>of bias | no serious<br>inconsistency | no serious indirectness | very<br>serious <sup>1</sup> | none                  | 3/16<br>(18.8%)      | 4/14<br>(28.6%) | RR<br>0.66<br>(0.18 to<br>2.44) | 97 fewer<br>per 1000<br>(from 234<br>fewer to<br>411<br>more) | LOW      | CRITICAL       |
| Disconti      | inuation (follo      | w-up mea                          | an 1 months; ass            | sessed with: Nu         | ımber of parti               | cipants lost to fol   | low-up)              |                 |                                 |   |          |                |
| 1             | randomised<br>trials | no<br>seriou<br>s risk<br>of bias | no serious inconsistency    | no serious indirectness | serious <sup>2</sup>         | none                  | 0/16<br>(0%)         | 0/15<br>(0%)    | not<br>pooled                   | not<br>pooled   | MODERATE | CRITICAL       |

CBT=cognitive behavioural therapy; CI=confidence interval; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standard mean difference; CPTS-RI=Child Post-Traumatic Stress-Reaction Index;

<sup>&</sup>lt;sup>1</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>2</sup> OIS not met (events<300)

Trauma-focused CBT versus waitlist, TAU or no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment                      |                              |                             |                            |                           |                       | No of pat                      | tients                                     | Effect                      |  |                 |                |
|----------------------|---------------------------------|------------------------------|-----------------------------|----------------------------|---------------------------|-----------------------|--------------------------------|--|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design                          | Risk of bias                 | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n           | Other consideration s | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality         | Importanc<br>e |
|                      | symptomatolog<br>ed by lower va |                              | d at endpoint (fo           | ollow-up 0.4-13            | weeks; measi              | ured with: SPTSS      | CPSS/CRI                       | ES/CRTI/UC                                 | LA PTSD-                    | RI/CPTS-RI   | change score;   | Better         |
| 13                   | randomised<br>trials            | serious <sup>1</sup>         | very serious <sup>2</sup>   | no serious<br>indirectness | no serious<br>imprecision | none                  | 440                            | 432  | -                           | SMD<br>1.21<br>lower<br>(1.59 to<br>0.83<br>lower)           | VERY LOW        | CRITICAL       |
|                      | symptomatologed by lower va     |                              | d at 1-3 month fo           | ollow-up (follow           | w-up 1-3 mont             | hs; measured wit      | h: IES/SPT                     | SS/CRIES/U                                 | CLA PTSE                    | -RI/CPTS-R   | I change score  | ; Better       |
| 5                    | randomised<br>trials            | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>      | none                  | 150                            | 151  | -                           | SMD<br>1.28<br>lower<br>(1.68 to<br>0.87<br>lower)           | LOW             | CRITICAL       |
| PTSD s               | ymptomatolog                    | gy self-rate                 | d at 6-month fol            | low-up (follow-            | up mean 6 mc              | onths; measured v     | with: SPTS                     | S change so                                | ore; Bette                  | r indicated l  | by lower values | s)             |
| 1                    | randomised<br>trials            | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup>      | none                  | 19                             | 20   | -                           | SMD<br>0.55<br>lower<br>(1.19<br>lower to<br>0.09<br>higher) | VERY LOW        | CRITICAL       |
| PTSD s               | ymptomatolog                    | gy self-rate                 | d at 12-18 month            | n follow-up (fol           | low-up 12-18 ı            | months; measure       | d with: CP                     | SS/SPTSS c                                 | hange sco                   | re; Better in  | dicated by low  | er values)     |
| 2                    | randomised                      | serious1                     | serious <sup>5</sup>        | no serious                 | serious <sup>3</sup>      | none                  | 55                             | 59   | _                           | SMD 0.6  |                 | CRITICAL       |

| Quality              | assessment                      |                                  |                             |                            |                           |                             | No of pat                      | tients                                     | Effect                      |  |                 |             |
|----------------------|---------------------------------|----------------------------------|-----------------------------|----------------------------|---------------------------|-----------------------------|--------------------------------|--|-----------------------------|--|-----------------|-------------|
| No of<br>studi<br>es | Design                          | Risk of<br>bias                  | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n           | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality         | Importance  |
|                      |                                 |                                  |                             |                            |                           |                             |                                |  |                             | (1.16 to<br>0.04<br>lower)                         |                 |             |
|                      | symptomatolog<br>ed by lower va |                                  | n-rated at endpoi           | nt (follow-up 8            | -20 weeks; me             | easured with: CAF           | PS/K-SADS                      | -E: PTSD/AI                                | DIS-C:PTS                   | D/CPTSDI; o  | change score;   | Better      |
| 7                    | randomised<br>trials            | no<br>serious<br>risk of<br>bias | very serious <sup>5</sup>   | no serious<br>indirectness | no serious<br>imprecision | none                        | 201                            | 208  | -                           | SMD<br>1.47<br>lower<br>(2.03 to<br>0.9<br>lower)  | LOW             | CRITICAL    |
|                      | symptomatolog                   |                                  | n-rated at 3-mont           | h follow-up (fo            | llow-up mean              | 3 months; measu             | ured with: (                   | CAPS/K-SAE                                 | S-E: PTSI                   | D/ADIS-C:P1  | rSD change sc   | ore; Better |
| 3                    | randomised<br>trials            | serious <sup>1</sup>             | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>      | reporting bias <sup>6</sup> | 59                             | 54   | -                           | SMD<br>0.75<br>lower<br>(1.14 to<br>0.37<br>lower) | VERY LOW        | CRITICAL    |
| PTSD s               | symptomatolog                   | gy cliniciar                     | n-rated at 6-mont           | h follow-up (fo            | llow-up mean              | 6 months; measu             | ured with: (                   | CAPS/K-SAE                                 | S-E: PTSI                   | D; Better inc                                      | licated by lowe | r values)   |
| 2                    | randomised<br>trials            | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>      | reporting bias <sup>6</sup> | 47                             | 42   | -                           | SMD<br>0.69<br>lower<br>(1.12 to<br>0.25<br>lower) | LOW             | CRITICAL    |

| Quality              | assessment                   |                                  |                             |                            |                      |                       | No of par                      | tients                                     | Effect                           |  |                 |                |
|----------------------|------------------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------|--------------------------------|--|----------------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design                       | Risk of<br>bias                  | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI)      | Absolute   | Quality         | Importanc<br>e |
| 2                    | randomised<br>trials         | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | reporting bias6       | 46                             | 42   | -                                | SMD<br>0.63<br>lower<br>(1.09 to<br>0.16<br>lower)           | LOW             | CRITICAL       |
| PTSD s               | ymptomatolog                 | gy cliniciar                     | -rated at 2-year            | follow-up (follo           | ow-up mean 2         | years; measured       | with: K-SA                     | DS-E: PTSE                                 | change s                         | core; Better   | indicated by le | ower values)   |
| 1                    | randomised<br>trials         | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness    | serious <sup>4</sup> | reporting bias6       | 21                             | 14   | -                                | SMD<br>0.22<br>lower<br>(0.9<br>lower to<br>0.46<br>higher)  | VERY LOW        | CRITICAL       |
| Remiss               | sion at endpoi               | nt (follow-ս                     | ıp 8-12 weeks; a            | ssessed with: I            | Number of peo        | ople no longer me     | eting diag                     | nostic criter                              | ia for PTS                       | D)   |                 |                |
| 5                    | randomised<br>trials         | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | none                  | 94/132<br>(71.2%)              | 59/145<br>(40.7%)                          | RR<br>1.75<br>(1.33 to<br>2.3)   | 305 more<br>per 1000<br>(from 134<br>more to<br>529<br>more) | MODERATE        | CRITICAL       |
|                      | sion at 1-3 mor<br>for PTSD) | nth follow-                      | up (follow-up 1-3           | months; asse               | ssed with: Nu        | mber of people no     | o longer ab                    | ove thresho                                | old on a sc                      | ale for PTSI   | or meeting di   | agnostic       |
| 2                    | randomised<br>trials         | no<br>serious<br>risk of<br>bias | no serious inconsistency    | no serious indirectness    | serious <sup>7</sup> | reporting bias6       | 31/46<br>(67.4%)               | 4/44<br>(9.1%)                             | RR<br>7.33<br>(2.84 to<br>18.91) | 575 more<br>per 1000<br>(from 167<br>more to                 | LOW             | CRITICAL       |

| Quality              | assessment           |                                  |                                  |                            |                              |                             | No of pat                      | ients                                      | Effect                           |  |                 |                |
|----------------------|----------------------|----------------------------------|----------------------------------|----------------------------|------------------------------|-----------------------------|--------------------------------|--|----------------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias                  | Inconsistenc<br>y                | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI)      | Absolute   | Quality         | Importanc<br>e |
|                      |                      |                                  |                                  |                            |                              |                             |                                |  |                                  | 1000<br>more)  |                 |                |
|                      | ion at 12-18 m       |                                  |                                  | 12-18 months;              | assessed with                | : Number of peop            | ole no long                    | er meeting o                               | diagnostic                       | criteria for l   | PTSD/scoring    | above          |
| 2                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency      | no serious<br>indirectness | serious4                     | reporting bias <sup>6</sup> | 42/108<br>(38.9%)              | 34/105<br>(32.4%)                          | RR<br>1.19<br>(0.85 to<br>1.67)  | 62 more<br>per 1000<br>(from 49<br>fewer to<br>217<br>more)  | LOW             | CRITICAL       |
|                      |                      |                                  | p 10-13 weeks; a improved' on Co |                            | Number of pe                 | ople showing clin           | ically sign                    | ificant impro                              | ovement, k                       | oased on reli  | iable change ir | ndices         |
| 3                    | randomised<br>trials | very<br>serious <sup>1</sup>     | serious <sup>5</sup>             | no serious<br>indirectness | serious <sup>7</sup>         | none                        | 58/101<br>(57.4%)              | 10/102<br>(9.8%)                           | RR<br>5.35<br>(1.64 to<br>17.39) | 426 more<br>per 1000<br>(from 63<br>more to<br>1000<br>more) | VERY LOW        | CRITICAL       |
| Anxiety              | symptoms at          | endpoint (                       | follow-up 2-20 w                 | veeks; measure             | ed with: HADS                | -A/SCARED/RCM               | AS/SCAS/                       | BAI change                                 | score; Bet                       | ter indicated  | d by lower valu | ies)           |
| 8                    | randomised<br>trials | very<br>serious <sup>1</sup>     | serious <sup>5</sup>             | no serious<br>indirectness | no serious<br>imprecision    | none                        | 268                            | 286  | -                                | SMD<br>0.81<br>lower<br>(1.23 to<br>0.4<br>lower)            | VERY LOW        | IMPORTA<br>NT  |
|                      |                      |                                  |                                  |                            |                              | ed with: HADS-A/I           |                                |  | Better ind                       |  | wer values)     |                |
| 2                    | randomised<br>trials | very<br>serious <sup>1</sup>     | serious <sup>5</sup>             | no serious indirectness    | very<br>serious <sup>8</sup> | none                        | 31                             | 32   | -                                | SMD<br>0.34  | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment           |                              |                             |                            |                        |                       | No of par                      | tients                                     | Effect                      |  |                 |               |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|------------------------|-----------------------|--------------------------------|--|-----------------------------|--|-----------------|---------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n        | Other consideration s | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality         | Importance    |
|                      |                      |                              |                             |                            |                        |                       |                                |  |                             | lower<br>(1.18<br>lower to<br>0.5<br>higher)       |                 |               |
| Anxiet               | y symptoms at        | 6-month f                    | ollow-up (follow-           | up mean 6 mo               | nths; measure          | ed with: HADS-A       | change sco                     | re; Better ir                              | dicated by                  | y lower valu                                       | es)             |               |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>   | none                  | 19                             | 20   | -                           | SMD<br>0.87<br>lower<br>(1.53 to<br>0.21<br>lower) | VERY LOW        | IMPORTA<br>NT |
| Anxiety              | y symptoms at        | 12-18 mor                    | nth follow-up (fo           | low-up 12-18 n             | nonths; measi          | ured with: HADS-      | A/SCARED                       | change sco                                 | re; Better                  | indicated by                                       | y lower values  | )             |
| 2                    | randomised<br>trials | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>   | none                  | 55                             | 59   | -                           | SMD<br>0.76<br>lower<br>(1.22 to<br>0.3<br>lower)  | LOW             | IMPORTA<br>NT |
| Depres               | sion symptom         | s at endpo                   | oint (follow-up 2-          | 20 weeks; mea              | sured with: H          | ADS-D/CES-D/CD        | I/MFQ/DSR                      | S/BDI chang                                | ge score; I                 | Better indica                                      | ited by lower v | alues)        |
| 13                   | randomised<br>trials | serious <sup>1</sup>         | serious <sup>5</sup>        | no serious indirectness    | no serious imprecision | none                  | 411                            | 423  | -                           | SMD<br>0.72<br>lower<br>(1.03 to<br>0.41<br>lower) | LOW             | IMPORTA<br>NT |

| Quality              | assessment           |                      |                             |                            |                      |                             | No of par                      | tients                                     | Effect                      |  |                  |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|--------------------------------|--|-----------------------------|--|------------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias      | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality          | Importanc<br>e |
| 7                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | reporting bias6             | 191                            | 188  | -                           | SMD<br>0.62<br>lower<br>(0.87 to<br>0.36<br>lower) | VERY LOW         | IMPORTA<br>NT  |
| Depres values)       |                      | s at 6-mor           | th follow-up (fol           | low-up mean 6              | months; mea          | sured with: HADS            | S-D/CDI/MII                    | NI:Depression                              | on change                   | score; Bette                                       | er indicated by  | lower          |
| 3                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | reporting bias6             | 66                             | 63   | -                           | SMD<br>0.48<br>lower<br>(0.84 to<br>0.13<br>lower) | VERY LOW         | IMPORTA<br>NT  |
| Depres<br>lower v    |                      | s at 12-18           | month follow-up             | (follow-up 12-             | 18 months; m         | easured with: HA            | DS-D/CDI/I                     | MINI:Depres                                | sion/MFQ                    | change sco   | re; Better indic | ated by        |
| 4                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | none                        | 101                            | 102  | -                           | SMD 0.5<br>lower<br>(0.78 to<br>0.22<br>lower)     | LOW              | IMPORTA<br>NT  |
| Depres               | sion symptom         | ıs at 2 year         | follow-up (follow           | w-up mean 2 ye             | ears; measure        | d with: CDI chang           | ge score; B                    | Better indica                              | ted by low                  | er values)   |                  |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | very<br>serious8     | reporting bias <sup>6</sup> | 21                             | 15   | -                           | SMD<br>0.17<br>lower<br>(0.83<br>lower to          | VERY LOW         | IMPORTA<br>NT  |

| Quality         | assessment                  |                              |                             |                            |                           |                             | No of pat                      | tients                                     | Effect                      |  |                  |                |
|-----------------|-----------------------------|------------------------------|-----------------------------|----------------------------|---------------------------|-----------------------------|--------------------------------|--|-----------------------------|--|------------------|----------------|
| No of studi es  | Design                      | Risk of bias                 | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n           | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality          | Importanc<br>e |
|                 |                             |                              |                             |                            |                           |                             |                                |  |                             | 0.5<br>higher)   |                  |                |
| Emotio          | nal and behav               | ioural prob                  | olems at endpoir            | it (follow-up 6-1          | l3 weeks; mea             | asured with: SDQ            | -A/CBCL cl                     | hange score                                | ; Better in                 | dicated by I   | ower values)     |                |
| 5               | randomised<br>trials        | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | no serious<br>imprecision | none                        | 245                            | 231  | -                           | SMD<br>0.58<br>lower<br>(0.79 to<br>0.36<br>lower)           | LOW              | IMPORTA<br>NT  |
| Emotio          | nal and beavio              | oural probl                  | ems at 18-month             | follow-up (foll            | ow-up mean 1              | l8 months; measu            | red with: 9                    | SDQ change                                 | score; Be                   | tter indicate  | ed by lower val  | ues)           |
| 1               | randomised<br>trials        | serious<br>1                 | no serious inconsistency    | no serious indirectness    | serious <sup>3</sup>      | none                        | 36                             | 39   | -                           | MD 2.83<br>lower<br>(4.79 to<br>0.87<br>lower)               | LOW              | IMPORTA<br>NT  |
| Emotion values) |                             | ioural prob                  | olems-Externaliz            | ing at endpoint            | (follow-up 12             | -20 weeks; meas             | ured with:                     | CBCL Exter                                 | nalizing ch                 | ange score   | ; Better indicat | ed by lower    |
| 3               | randomised<br>trials        | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup>      | none                        | 105                            | 105  | -                           | SMD<br>0.25<br>lower<br>(0.67<br>lower to<br>0.16<br>higher) | VERY LOW         | IMPORTA<br>NT  |
|                 | nal and behaved by lower va |                              | olems-Externaliz            | ing at 3-month             | follow-up (fol            | low-up mean 3 m             | onths; mea                     | sured with:                                | CBCL Ext                    | ernalizing c   | hange score; E   | Better         |
| 2               | randomised trials           | serious <sup>1</sup>         | no serious inconsistency    | no serious indirectness    | serious <sup>3</sup>      | reporting bias <sup>6</sup> | 32                             | 24   | -                           | SMD<br>0.77  | VERY LOW         | IMPORTA<br>NT  |

| Quality              | assessment                      |                      |                             |                            |                      |                             | No of par                      | tients                                     | Effect                      |  |                 |                |
|----------------------|---------------------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|--------------------------------|--|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design                          | Risk of<br>bias      | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality         | Importanc<br>e |
|                      |                                 |                      |                             |                            |                      |                             |                                |  |                             | lower<br>(1.32 to<br>0.21<br>lower)                      |                 |                |
|                      | nal and behav<br>ed by lower va |                      | olems-Externaliz            | ing at 6-month             | follow-up (fol       | low-up mean 6 m             | onths; mea                     | sured with:                                | CBCL Ext                    | ernalizing c   | hange score; E  | Better         |
| 1                    | randomised<br>trials            | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | reporting bias <sup>6</sup> | 20                             | 12   | -                           | SMD<br>0.82<br>lower<br>(1.57 to<br>0.07<br>lower)       | VERY LOW        | IMPORTA<br>NT  |
|                      | nal and behaved by lower va     |                      | olems-Externaliz            | ing at 12-mont             | h follow-up (fo      | ollow-up mean 12            | months; m                      | neasured wit                               | h: CBCL E                   | Externalizing  | ı change score  | ; Better       |
| 1                    | randomised<br>trials            | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup> | reporting bias <sup>6</sup> | 20                             | 12   | -                           | SMD 0.7<br>lower<br>(1.44<br>lower to<br>0.04<br>higher) | VERY LOW        | IMPORTA<br>NT  |
|                      | nal and behaver values)         | ioural prob          | olems-Externaliz            | ing at 2-year fo           | llow-up (follo       | w-up mean 2 year            | s; measure                     | ed with: CB0                               | CL Externa                  | llizing chang  | ge score; Bette | r indicated    |
| 1                    | randomised<br>trials            | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | reporting bias <sup>6</sup> | 20                             | 12   | -                           | SMD<br>1.41<br>lower<br>(2.22 to<br>0.61<br>lower)       | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment           |                              |                             |                            |                      |                             | No of pat                      | ients                                      | Effect                      |  |                 |                |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|--------------------------------|--|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute   | Quality         | Importanc<br>e |
| Emotio<br>values)    |                      | ioural prob                  | lems-Internalizi            | ng at endpoint             | (follow-up 12-       | -20 weeks; measu            | red with: C                    | BCL Interna                                | alizing cha                 | nge score;   | Better indicate | d by lower     |
| 2                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | none                        | 85                             | 93   | -                           | SMD<br>0.61<br>lower<br>(1.03 to<br>0.2<br>lower)            | VERY LOW        | IMPORTA<br>NT  |
|                      | nal and behav        |                              | olems-Internalizi           | ng at 3-month t            | follow-up (foll      | ow-up mean 3 mo             | onths; meas                    | sured with:                                | CBCL Inte                   | rnalizing ch   | ange score; Be  | etter          |
| 1                    | randomised<br>trials | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup> | reporting bias <sup>6</sup> | 12                             | 12   | -                           | SMD<br>0.71<br>lower<br>(1.54<br>lower to<br>0.12<br>higher) | VERY LOW        | IMPORTA<br>NT  |
|                      |                      | -up 6-12 we                  | eks; measured               | with: KIDSCRE              |                      | HRQoL T-scores              |                                |  | etter indica                |  | er values)      | 1              |
| 2                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup> | none                        | 118                            | 101  | -                           | SMD<br>0.33<br>higher<br>(0.06 to<br>0.6<br>higher)          | VERY LOW        | IMPORTA<br>NT  |
|                      | _                    |                              |                             |                            |                      | CAPS: Functional            |                                |  | change s                    |  | indicated by lo |                |
| 2                    | randomised trials    | very<br>serious <sup>1</sup> | very serious <sup>2</sup>   | no serious indirectness    | serious <sup>4</sup> | none                        | 47                             | 48   | -                           | SMD<br>1.56  | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                            |                      |                             | No of pa                       | tients                                     | Effect                      |   |                  |               |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|--------------------------------|--|-----------------------------|---|------------------|---------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias      | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute  | Quality          | Importanc     |
|                      |                      |                      |                             |                            |                      |                             |                                |  |                             | (3.14<br>lower to<br>0.02<br>higher)                        |                  |               |
| Function values)     |                      | nt at 3-mor          | nth follow-up (fol          | low-up mean 3              | months; mea          | sured with: CAPS            | : Function                     | al impairme                                | nt; change                  | e score; Bet  | ter indicated by | y lower       |
| 2                    | randomised<br>trials | serious <sup>1</sup> | serious <sup>5</sup>        | no serious<br>indirectness | serious <sup>3</sup> | reporting bias <sup>6</sup> | 110                            | 110  | -                           | SMD<br>0.96<br>lower<br>(1.24 to<br>0.68<br>lower)          | VERY LOW         | IMPORTA<br>NT |
| Function values)     |                      | nt at 6-mor          | nth follow-up (fol          | low-up mean 6              | months; mea          | sured with: CAPS            | : Function                     | al impairme                                | nt; change                  | e score; Bet  | ter indicated by | y lower       |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup> | reporting bias <sup>6</sup> | 26                             | 28   | -                           | SMD<br>0.45<br>lower<br>(0.99<br>lower to<br>0.1<br>higher) | VERY LOW         | IMPORTA<br>NT |
| Function values)     | •                    | nt at 12-mo          | onth follow-up (fo          | ollow-up mean              | 12 months; m         | easured with: CA            | PS: Functi                     | onal impairr                               | nent; char                  | nge score; B  | etter indicated  | by lower      |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious<br>indirectness | serious <sup>3</sup> | reporting bias <sup>6</sup> | 25                             | 28   | -                           | SMD<br>1.28<br>lower<br>(1.88 to                            | VERY LOW         | IMPORTA<br>NT |

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of pat                      | tients                                     | Effect                      |   |          |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|--------------------------------|--|-----------------------------|---|----------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Waitlist,<br>TAU or<br>no<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute  | Quality  | Importanc<br>e |
|                      |                      |                                  |                             |                            |                      |                             |                                |  |                             | 0.69<br>lower)  |          |                |
| Global               | functioning at       | endpoint (                       | follow-up 10-20             | weeks; measur              | ed with: CGA         | S/fCPSS/GAF cha             | nge score                      | ; Better indi                              | cated by h                  | igher values  | s)       |                |
| 4                    | randomised<br>trials | very<br>serious <sup>1</sup>     | serious <sup>5</sup>        | no serious<br>indirectness | serious <sup>3</sup> | none                        | 153                            | 168  | -                           | SMD<br>1.25<br>higher<br>(0.65 to<br>1.85<br>higher)      | VERY LOW | IMPORTA<br>NT  |
| Global               | functioning at       | 3-month fo                       | ollow-up (follow-           | up mean 3 moi              | nths; measure        | d with: GAF; cha            | nge score;                     | Better indic                               | ated by hi                  | gher values   | )        |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness    | serious <sup>3</sup> | reporting bias <sup>6</sup> | 12                             | 12   | -                           | SMD<br>1.35<br>higher<br>(0.45 to<br>2.25<br>higher)      | VERY LOW | IMPORTA<br>NT  |
| Global               | functioning at       | 18-month                         | follow-up (follov           | /-up mean 18 m             | nonths; measu        | red with: CPSS o            | hange sco                      | re; Better in                              | dicated by                  | y higher valu   | ues)     |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup> | none                        | 36                             | 39   | -                           | SMD 0.1<br>higher<br>(0.35<br>lower to<br>0.56<br>higher) | LOW      | IMPORTA<br>NT  |
| Discont              | tinuation (follo     | w-up 0.4-2                       | 0 weeks; assess             | ed with: Numb              | er of participa      | nts lost to follow          | -up for any                    | / reason)                                  |                             |   |          |                |
| 18                   | randomised trials    | no<br>serious<br>risk of<br>bias | no serious inconsistency    | no serious indirectness    | serious <sup>9</sup> | none                        | 70/642<br>(10.9%)              | 46/613<br>(7.5%)                           | RR 1.3<br>(0.93 to<br>1.83) | 23 more<br>per 1000<br>(from 5                            | MODERATE | CRITICAL       |

| Quality     | assessment |              |                   |                  |                 |                       | No of pat             | tients                                | Effect                      |                      |         |           |
|-------------|------------|--------------|-------------------|------------------|-----------------|-----------------------|-----------------------|---------------------------------------|-----------------------------|----------------------|---------|-----------|
| No of studi | Design     | Risk of bias | Inconsistenc<br>y | Indirectnes<br>s | Imprecisio<br>n | Other consideration s | Trauma - focuse d CBT | Waitlist,<br>TAU or<br>no<br>treatmen | Relativ<br>e<br>(95%<br>CI) | Absolute             |         | Importanc |
|             |            |              |                   |                  |                 |                       |                       | t                                     |                             |                      | Quality | е         |
|             |            |              |                   |                  |                 |                       |                       |                                       |                             | fewer to<br>62 more) |         |           |

ADIS-C=Anxiety Disorder Interview Schedule-Child version: BAI=Beck Anxiety Index; BDI=Beck Depression Inventory; CAPS=Clinician Administered PTSD Symptom; CBCL=Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI=Children's Depression Inventory; CES-D=Centre for Epidemiological Studies-Depression; CGAS= Children's Global Assessment Scale; CGI=Clinical Global Impression; CI=confidence interval; CPSS=Child PTSD Symptom Scale; CPTS-RI=Child Post-Traumatic Symptom-Reaction Index; CRIES=Children's Revised Impact of Event Scale; CRTI=Children's Response to Trauma Inventory; DSRS=Depression Self-Rating Scale; GAF=Global Assessment of Functioning; HRQoL=Health-Related Quality of Life; KIDSCREEN-27=Health-related quality of life questionnaire for children, young people and their parents; K-SADS-E=Kiddie Schedule for Affective Disorders and Schizophrenia-Epidemiological; HADS-A/D=Hospital Anxiety and Depression Scale-Anxiety/Depression; ILK=an instrument to measure quality of life in children and adolescents; MFQ=Mood and Feeling Questionnaire; PTSD=post-traumatic stress disorder; RCMAS=Revised Children's Manifest Anxiety Scale; RR=risk ratio; SAS-SR=Social Adjustment Scale-Self Report; SCARED=Screen for Child Anxiety Related Disorders; SCAS=Spence Children's Anxiety Scale; SDQ =Strength and Difficulties Questionnaires; SMD=standard mean difference; SPTSS=Screen for Post-Traumatic Stress Symptoms; TAU=treatment as usual: UCLA PTSD-Reaction Index

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> Considerable heterogeneity (I2>80%)

<sup>&</sup>lt;sup>3</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>5</sup> Substantial heterogeneity (I2=>50%)

<sup>&</sup>lt;sup>6</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>7</sup> OIS not met (events<300)

<sup>8 95%</sup> CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>9</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

Trauma-focused CBT versus supportive counselling for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                              |                             |                            |                      |                       | No of pat             | tients                            | Effect                      |  |                  |            |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------|-----------------------|-----------------------------------|-----------------------------|--|------------------|------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e                                       | Quality          | Importance |
| PTSD s<br>lower v    | •                    | gy self-rate                 | ed at endpoint (fo          | ollow-up 6-15 v            | veeks; measu         | red with: CRIES/T     | SCC-PTSE              | O/UCLA PTSD                       | -RI/CPSS                    | change sco   | re; Better indic | ated by    |
| 5                    | randomised<br>trials | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 165                   | 160                               | -                           | SMD<br>0.49<br>lower<br>(0.71 to<br>0.26<br>lower) | LOW              | CRITICAL   |
| PTSD s               | symptomatolog        | gy self-rate                 | ed at 3-month fol           | low-up (follow             | -up mean 3 m         | onths; measured       | with: CRIE            | S change sco                      | re; Better                  | indicated b  | y lower values   | )          |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 10                    | 10                                | -                           | SMD<br>1.58<br>lower<br>(2.62 to<br>0.55<br>lower) | VERY LOW         | CRITICAL   |
| PTSD s               |                      | gy self-rate                 | ed at 6-month fol           | low-up (follow             | -up mean 6 m         | onths; measured       | with: TSC             | C-PTSD/VCPS                       | S change                    | score; Bett  | er indicated by  | lower      |
| 2                    | randomised<br>trials | serious <sup>1</sup>         | serious3                    | no serious<br>indirectness | serious <sup>2</sup> | reporting bias4       | 60                    | 60                                | -                           | SMD 0.7<br>lower<br>(1.29 to<br>0.11<br>lower)     | VERY LOW         | CRITICAL   |
| PTSD s               | •                    | gy self-rate                 | ed at 12-17 mont            | h follow-up (fo            | llow-up 12-17        | months; measure       | ed with: TS           | CC-PTSD/CP                        | SS change                   | e score; Bet                                       | ter indicated b  | y lower    |
| 3                    | randomised<br>trials | serious <sup>1</sup>         | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias4       | 91                    | 90                                | -                           | SMD<br>0.69<br>lower<br>(0.99 to                   | VERY LOW         | CRITICAL   |

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of pat             | ients                             | Effect                      |  |                  |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|--|------------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Quality          | Importanc<br>e |
|                      |                      |                                  |                             |                            |                      |                             |                       |                                   |                             | 0.39<br>lower)   |                  |                |
| PTSD s<br>values)    |                      | gy cliniciar                     | n-rated at endpo            | int (follow-up 8           | 3-14 weeks; m        | easured with: K-S           | ADS-PL: F             | PTSD/CPSS-I/                      | CAPS; cha                   | ange score   | Better indicate  | ed by lower    |
| 3                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | none                        | 121                   | 110                               | -                           | SMD<br>0.71<br>lower<br>(1.1 to<br>0.31<br>lower)            | MODERATE         | CRITICAL       |
| PTSD s               | ymptomatolog         | gy cliniciar                     | n-rated at 3-mon            | th follow-up (fo           | ollow-up mear        | 3 months; meas              | ured with:            | CAPS change                       | e score; B                  | etter indica   | ted by lower va  | lues)          |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious indirectness    | serious <sup>5</sup> | reporting bias <sup>4</sup> | 26                    | 24                                | -                           | SMD<br>0.25<br>lower<br>(0.81<br>lower to<br>0.31<br>higher) | LOW              | CRITICAL       |
| PTSD s               | ymptomatolog         | gy cliniciar                     | n-rated at 6-mon            | th follow-up (fo           | ollow-up mear        | 6 months; meas              | ured with:            | CAPS change                       | e score; B                  | etter indica   | ted by lower va  | lues)          |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup> | reporting bias <sup>4</sup> | 26                    | 23                                | -                           | SMD<br>0.43<br>lower (1<br>lower to<br>0.13<br>higher)       | LOW              | CRITICAL       |
| PTSD s<br>values)    |                      | gy cliniciar                     | n-rated at 12-mor           | nth follow-up (            | follow-up mea        | in 12 months; me            | asured wit            | h: CAPS/CPS                       | S-I chang                   | e score; Be  | tter indicated b | y lower        |
| 2                    | randomised<br>trials | no<br>serious                    | no serious inconsistency    | no serious<br>indirectness | serious <sup>2</sup> | reporting bias4             | 56                    | 53                                | -                           | SMD<br>0.89  | LOW              | CRITICAL       |

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of pat             | tients                            | Effect                          |   |          |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|---------------------------------|---|----------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI)     | Absolut<br>e  | Quality  | Importane<br>e |
|                      |                      | risk of<br>bias                  |                             |                            |                      |                             |                       |                                   |                                 | lower<br>(1.28 to<br>0.49<br>lower)                             | _        |                |
| Remiss               | sion at endpoi       | nt (follow-u                     | up 8-15 weeks; a            | ssessed with:              | Number of pe         | ople no longer m            | eeting diag           | nostic criteria                   | a for PTSE                      | ))  |          |                |
| 4                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | none                        | 71/115<br>(61.7%)     | 35/93<br>(37.6%)                  | RR<br>1.67<br>(1.25 to<br>2.23) | more per<br>1000<br>(from 94<br>more to<br>463<br>more)         | MODERATE | CRITICAL       |
| Remiss               | sion at 6-mont       | h follow-uլ                      | (follow-up mea              | n 6 months; as             | sessed with:         | Number of people            | e no longe            | r meeting diag                    | gnostic cri                     | teria for PT  | SD)      |                |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | none                        | 12/19<br>(63.2%)      | 5/19<br>(26.3%)                   | RR 2.4<br>(1.05 to<br>5.49)     | 368<br>more per<br>1000<br>(from 13<br>more to<br>1000<br>more) | MODERATE | CRITICAL       |
| Remiss               | sion at 12-mon       | th follow-u                      | ip (follow-up me            | an 12 months;              | assessed wit         | h: Number of peo            | ple no long           | ger meeting d                     | iagnostic                       | criteria for  | PTSD)    |                |
| 2                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>4</sup> | 46/60<br>(76.7%)      | 29/58<br>(50%)                    | RR<br>1.56<br>(1.17 to<br>2.08) | 280<br>more per<br>1000<br>(from 85<br>more to<br>540<br>more)  | LOW      | CRITICAL       |

| Quality     | assessment           |                                  |                             |                            |                      |                             | No of pat             | tients                            | Effect                          |   |                |                |
|-------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|---------------------------------|---|----------------|----------------|
| No of studi | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI)     | Absolut<br>e  | Quality        | Importanc<br>e |
| 1           | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>4</sup> | 23/31<br>(74.2%)      | 8/30<br>(26.7%)                   | RR<br>2.78<br>(1.48 to<br>5.22) | 475<br>more per<br>1000<br>(from<br>128<br>more to<br>1000<br>more) | LOW            | CRITICAL       |
|             |                      |                                  |                             |                            |                      | : Number of peop            |                       |                                   |                                 |   | t (based on R0 |                |
| 1           | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>4</sup> | 22/31<br>(71%)        | 12/30<br>(40%)                    | RR<br>1.77<br>(1.08 to<br>2.9)  | 308<br>more per<br>1000<br>(from 32<br>more to<br>760<br>more)      | LOW            | CRITICAL       |
| Dissoc      | iative symptor       | ns at endp                       | oint (follow-up r           | nean 12 weeks              | ; measured w         | ith: TSCC-Dissoc            | iation char           | nge score; Be                     | tter indica                     | ted by lowe   | r values)      |                |
| 1           | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup> | reporting bias <sup>4</sup> | 41                    | 41                                | -                               | SMD<br>0.27<br>lower<br>(0.71<br>lower to<br>0.16<br>higher)        | VERY LOW       | IMPORTA<br>NT  |
| Dissoc      | iative symptor       | ns at 6-mo                       | onth follow-up (fo          | ollow-up mean              | 6 months; me         | asured with: TSC            | C-Dissocia            | ation change                      | score; Bet                      | tter indicate   | d by lower val | ues)           |
| 1           | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>4</sup> | 41                    | 41                                | -                               | SMD 0.7<br>lower<br>(1.15 to  | VERY LOW       | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                            |                           |                             | No of pat             | tients                            | Effect                      |  | <b>,</b>        |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|---------------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias      | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n           | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Quality         | Importanc<br>e |
|                      |                      |                      |                             |                            |                           |                             |                       |                                   |                             | 0.25<br>lower)   |                 |                |
| Dissoc               | iative symptor       | ns at 12-m           | onth follow-up (            | follow-up meai             | n 12 months; r            | measured with: T            | SCC-Disso             | ciation chang                     | je score; E                 | Better indic   | ated by lower v | alues)         |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>      | reporting bias <sup>4</sup> | 41                    | 41                                | -                           | SMD<br>0.49<br>lower<br>(0.93 to<br>0.05<br>lower)       | VERY LOW        | IMPORTA<br>NT  |
| Anxiety              | y symptoms at        | endpoint             | (follow-up 8-12 v           | veeks; measur              | ed with: STAI-            | -State/SCARED/T             | SCC:Anxie             | ty change sc                      | ore; Bette                  | r indicated  | by lower values | s)             |
| 4                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | no serious<br>imprecision | reporting bias <sup>4</sup> | 222                   | 211                               | -                           | SMD<br>0.29<br>lower<br>(0.48 to<br>0.1<br>lower)        | LOW             | IMPORTA<br>NT  |
| Anxiety              | y symptoms at        | 6-month f            | ollow-up (follow            | -up mean 6 mc              | onths; measur             | ed with: STAI-Sta           | te change             | score; Better                     | indicated                   | by lower va  | alues)          |                |
| 2                    | randomised<br>trials | serious <sup>1</sup> | serious <sup>3</sup>        | no serious<br>indirectness | serious <sup>5</sup>      | reporting bias <sup>4</sup> | 121                   | 112                               | -                           | SMD 0.3<br>lower<br>(0.82<br>lower to<br>0.22<br>higher) | VERY LOW        | IMPORTA<br>NT  |
| Anxiety              | y symptoms at        | 12-month             | follow-up (follow           | w-up mean 12               | months; meas              | ured with: STAI-S           | State chang           | ge score; Bett                    | er indicat                  | ed by lower  | values)         |                |
| 2                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>5</sup>      | reporting bias <sup>4</sup> | 123                   | 114                               | -                           | SMD<br>0.17<br>lower<br>(0.51<br>lower to                | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                           |                            |                              |                             | No of pat             | tients                            | Effect                      |  |                 |                |
|----------------------|----------------------|----------------------|---------------------------|----------------------------|------------------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias         | Inconsistenc<br>y         | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Quality         | Importanc<br>e |
|                      |                      |                      |                           |                            |                              |                             |                       |                                   |                             | 0.17<br>higher)  |                 |                |
| Depres               | sion symptom         | s at endpo           | oint (follow-up 6-        | -15 weeks; mea             | asured with: B               | DI/CES-D/CDI/TS             | CC:Depres             | sion change                       | score; Be                   | tter indicate  | ed by lower val | ues)           |
| 7                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency  | no serious<br>indirectness | no serious<br>imprecision    | reporting bias <sup>4</sup> | 282                   | 270                               | -                           | SMD<br>0.41<br>lower<br>(0.67 to<br>0.16<br>lower)           | LOW             | IMPORTA<br>NT  |
| Depres               | sion symptom         | ıs at 3-mor          | nth follow-up (fo         | llow-up mean 3             | 3 months; mea                | asured with: CES            | D/MINI:De             | pression chai                     | nge score                   | Better indi  | icated by lower | values)        |
| 2                    | randomised<br>trials | serious <sup>1</sup> | very serious <sup>7</sup> | no serious<br>indirectness | very<br>serious <sup>8</sup> | reporting bias <sup>4</sup> | 36                    | 34                                | -                           | SMD<br>0.46<br>lower<br>(2.26<br>lower to<br>1.33<br>higher) | VERY LOW        | IMPORTA<br>NT  |
| Depres               | sion symptom         | s at 6-mor           | nth follow-up (fo         | llow-up mean (             | 6 months; mea                | asured with: BDI/0          | CDI/MINI:D            | epression cha                     | ange scor                   | e; Better in   | dicated by low  | er values)     |
| 4                    | randomised<br>trials | serious <sup>1</sup> |                           | no serious<br>indirectness | serious <sup>5</sup>         | reporting bias <sup>4</sup> | 166                   | 154                               | -                           | SMD 0.3<br>lower<br>(0.74<br>lower to<br>0.13<br>higher)     | VERY LOW        | IMPORTA<br>NT  |
| Depres               | sion symptom         | s at 12-17           | month follow-up           | (follow-up 12              | -17 months; m                | neasured with: BD           | I/CDI/MINI            | :Depression of                    | change sc                   | ore; Better  | indicated by lo | wer values)    |
| 5                    | randomised<br>trials | serious <sup>1</sup> | serious <sup>3</sup>      | no serious indirectness    | serious <sup>5</sup>         | reporting bias <sup>4</sup> | 198                   | 186                               | -                           | SMD<br>0.34<br>lower<br>(0.74                                | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment                   |                      |                             |                            |                      |                             | No of pat             | ients                             | Effect                      |   |                  |                |
|----------------------|------------------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|---|------------------|----------------|
| No of<br>studi<br>es | Design                       | Risk of bias         | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e  | Quality          | Importanc<br>e |
|                      |                              |                      |                             |                            |                      |                             |                       |                                   |                             | lower to<br>0.07<br>higher)                                   |                  |                |
| Emotion lower v      |                              | ioural prol          | olems-Internalizi           | ng at endpoint             | (follow-up m         | ean 12 weeks; me            | asured wit            | h: CBCL Inte                      | rnalizing o                 | hange sco   | re; Better indic | ated by        |
| 2                    | randomised<br>trials         | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 129                   | 132                               | -                           | SMD<br>0.08<br>lower<br>(0.33<br>lower to<br>0.16<br>higher)  | VERY LOW         | IMPORTA<br>NT  |
|                      | nal and behaved by lower va  |                      | olems-Internalizi           | ng at 6-month              | follow-up (fol       | low-up mean 6 m             | onths; mea            | sured with: C                     | BCL Inter                   | nalizing ch   | ange score; Be   | etter          |
| 2                    | randomised<br>trials         | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>9</sup> | reporting bias <sup>4</sup> | 116                   | 108                               | -                           | SMD<br>0.17<br>higher<br>(0.19<br>lower to<br>0.53<br>higher) | VERY LOW         | IMPORTA<br>NT  |
|                      | onal and behaved by lower va |                      | olems-Internalizi           | ng at 12-mont              | h follow-up (fo      | ollow-up mean 12            | months; m             | easured with                      | : CBCL In                   | ternalizing   | change score;    | Better         |
| 2                    | randomised<br>trials         | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 119                   | 109                               | -                           | SMD<br>0.02<br>higher<br>(0.24<br>lower to                    | VERY LOW         | IMPORTA<br>NT  |

| Quality              | assessment                  |                      |                             |                            |                      |                             | No of pat             | tients                            | Effect                      |   |                 |                |
|----------------------|-----------------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|---|-----------------|----------------|
| No of<br>studi<br>es | Design                      | Risk of bias         | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e  | Quality         | Importanc<br>e |
|                      |                             |                      |                             |                            |                      |                             |                       |                                   |                             | 0.28<br>higher)   |                 |                |
| Emotio<br>lower v    |                             | ioural prol          | olems-Externaliz            | ing at endpoin             | t (follow-up m       | nean 12 months; r           | neasured v            | vith: CBCL Ex                     | cternalizin                 | g change s  | core; Better in | dicated by     |
| 2                    | randomised<br>trials        | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 129                   | 132                               | -                           | SMD<br>0.15<br>lower<br>(0.4<br>lower to<br>0.09<br>higher)   | VERY LOW        | IMPORTA<br>NT  |
|                      | nal and behaved by lower va |                      | olems-Externaliz            | ing at 6-month             | follow-up (fo        | llow-up mean 6 m            | onths; me             | asured with:                      | CBCL Ext                    | ernalizing c  | hange score; E  | Better         |
| 2                    | randomised<br>trials        | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 116                   | 108                               | -                           | SMD<br>0.04<br>higher<br>(0.22<br>lower to<br>0.31<br>higher) | VERY LOW        | IMPORTA<br>NT  |
|                      | nal and behaved by lower va |                      | olems-Externaliz            | ing at 12-mont             | th follow-up (f      | ollow-up mean 12            | months; r             | neasured with                     | h: CBCL E                   | xternalizing  | g change score  | ; Better       |
| 2                    | randomised<br>trials        | serious <sup>1</sup> | serious <sup>3</sup>        | no serious<br>indirectness | serious <sup>9</sup> | reporting bias <sup>4</sup> | 119                   | 109                               | -                           | SMD<br>0.18<br>higher<br>(0.27<br>lower to<br>0.62<br>higher) | VERY LOW        | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                            |                      |                             | No of pat             | tients                            | Effect                      |   |                  |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|---|------------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias         | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e  | Quality          | Importanc<br>e |
| Behavi               | our problems         | at endpoin           | t (follow-up 8-12           | weeks; meas                | ured with: CB        | CL total score; ch          | ange score            | e; Better indic                   | ated by lo                  | wer values  | )                |                |
| 3                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 193                   | 192                               | -                           | SMD<br>0.11<br>lower<br>(0.31<br>lower to<br>0.09<br>higher)  | VERY LOW         | IMPORTA<br>NT  |
| Behavi               | our problems         | at 6-month           | follow-up (follo            | w-up mean 6 n              | nonths; meas         | ured with: CBCL t           | otal score            | ; change scor                     | e; Better i                 | ndicated by   | / lower values)  |                |
| 2                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 116                   | 108                               | -                           | SMD<br>0.08<br>higher<br>(0.18<br>lower to<br>0.34<br>higher) | VERY LOW         | IMPORTA<br>NT  |
| Behavi               | our problems         | at 12-mont           | th follow-up (foll          | ow-up mean 1               | 2 months; mea        | asured with: CBC            | L total sco           | re; change so                     | ore; Bette                  | r indicated   | by lower value   | s)             |
| 2                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>4</sup> | 119                   | 109                               | -                           | SMD<br>0.04<br>higher<br>(0.32<br>lower to<br>0.41<br>higher) | VERY LOW         | IMPORTA<br>NT  |
| Function values      | •                    | nt at 3-mor          | nth follow-up (fo           | llow-up mean 3             | 3 months; mea        | asured with: CAP            | S: Function           | nal impairmer                     | ıt; change                  | score; Bet  | ter indicated by | lower          |
| 1                    | randomised trials    | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>5</sup> | reporting bias <sup>4</sup> | 26                    | 24                                | -                           | SMD<br>0.43<br>lower (1                                       | VERY LOW         | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                            |                              |                             | No of pat             | ients                             | Effect                      |   |                  |               |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|-----------------------|-----------------------------------|-----------------------------|---|------------------|---------------|
| No of<br>studi<br>es | Design               | Risk of bias         | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e  | Quality          | Importance    |
|                      |                      |                      |                             |                            |                              |                             |                       |                                   |                             | lower to<br>0.13<br>higher)                                   |                  |               |
| Function values      |                      | nt at 6-mor          | nth follow-up (fo           | llow-up mean (             | 6 months; mea                | asured with: CAP            | S: Function           | nal impairmer                     | nt; change                  | score; Bet  | ter indicated by | y lower       |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>8</sup> | reporting bias <sup>4</sup> | 26                    | 23                                | -                           | SMD<br>0.01<br>higher<br>(0.55<br>lower to<br>0.57<br>higher) | VERY LOW         | IMPORTA<br>NT |
| Function values      |                      | nt at 12-mo          | onth follow-up (f           | ollow-up mean              | 12 months; m                 | neasured with: CA           | APS: Funct            | ional impairm                     | ent; chan                   | ge score; B   | etter indicated  | by lower      |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | reporting bias <sup>4</sup> | 25                    | 23                                | -                           | SMD<br>1.12<br>lower<br>(1.73 to<br>0.5<br>lower)             | VERY LOW         | IMPORTA<br>NT |
| Global               | functioning at       | endpoint             | (follow-up 14-15            | weeks; measu               | red with: CGA                | S; change score             | Better ind            | icated by hig                     | her values                  | s)  |                  |               |
| 2                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>         | reporting bias <sup>4</sup> | 50                    | 49                                | -                           | SMD<br>1.08<br>higher<br>(0.65 to<br>1.5                      | VERY LOW         | IMPORTA<br>NT |

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of pat             | ients                             | Effect                          |   |          |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|-----------------------------------|---------------------------------|---|----------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Supportiv<br>e<br>counsellin<br>g | Relativ<br>e<br>(95%<br>CI)     | Absolut<br>e  | Quality  | Importanc<br>e |
| 1                    | randomised<br>trials | serious<br>1                     | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                        | 19                    | 19                                | -                               | SMD<br>1.05<br>higher<br>(0.37 to<br>1.73<br>higher)        | LOW      | IMPORTA<br>NT  |
| Global               | functioning at       | 12-month                         | follow-up (follow           | w-up mean 12 ı             | months; meas         | ured with: CGAS             | change so             | core; Better ir                   | ndicated b                      | y higher va   | lues)    |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>4</sup> | 31                    | 30                                | -                               | SMD 1<br>higher<br>(0.47 to<br>1.54<br>higher)              | VERY LOW | IMPORTA<br>NT  |
| Discon               | tinuation (follo     | ow-up 3-15                       | weeks; assesse              | ed with: Numbe             | er of participa      | nts lost to follow-         | up for any            | reason)                           |                                 |   |          |                |
| 8                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup> | none                        | 80/347<br>(23.1%)     | 95/331<br>(28.7%)                 | RR<br>0.78<br>(0.61 to<br>1.01) | 63 fewer<br>per 1000<br>(from<br>112<br>fewer to<br>3 more) | MODERATE | CRITICAL       |

BDI= Beck Depression Inventory; CAPS= Clinician Administered PTSD Symptom; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CES-D= Centre for Epidemiological Studies-Depression; CGAS= Children's Global Assessment Scale; CI=confidence interval; CPSS= Child PTSD Symptom Scale; CRIES= Children's Revised Impact of Event Scale; K-SADS= Kiddie Schedule for Affective Disorders and Schizophrenia-Epidemiological; PTSD=post-traumatic stress disorder; RCI=Reliable Change Indecies; RR=risk ratio; SCARED=Screen for Child Anxiety Related Disorders; SMD=standardised mean difference; STAI=State-Trait Anxiety Inventory; TSCC = Trauma Symptom Checklist for Children; UCLA PTSD-RI=UCLA PTSD-Reaction Index

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains <sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Substantial heterogeneity (I2>50%)

<sup>&</sup>lt;sup>4</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>5</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>6</sup> OIS not met (events<300)

Trauma-focused CBT versus eye movement desensitisation and reprocessing (EMDR) for the delayed treatment (>3 months) of clinically important symptoms/PTSD - Single incident index trauma

| Quality              | assessment           |                              |                             |                            |                      |                             | No of pat             | ients  | Effect                      |  |             |                |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Eye movement<br>desensitisation<br>and<br>reprocessing<br>(EMDR) | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Qualit<br>y | Importanc<br>e |
| PTSD s               | ymptomatolog         | gy self-rate                 | ed at endpoint (fo          | llow-up mean               | 6 weeks; mea         | sured with: CRTI            | change sc             | ore; Better indicate   | d by lowe                   | r values)  |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.13<br>lower<br>(0.56<br>lower to<br>0.29<br>higher) | VERY<br>LOW | CRITICAL       |
| PTSD s               | ymptomatolog         | gy self-rate                 | ed at 3-month fol           | low-up (follow-            | up mean 3 mo         | onths; measured             | with: CRTI            | change score; Bet  | ter indicat                 | ed by lower  | values)     |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.35<br>lower<br>(0.77<br>lower to<br>0.08<br>higher) | VERY<br>LOW | CRITICAL       |
| PTSD s               | ymptomatolog         | gy self-rate                 | ed at 12-month fo           | llow-up (follow            | -up mean 12          | months; measure             | d with: CR            | TI change score; B   | etter indic                 | ated by low  | er value    | s)             |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.24<br>lower   | VERY<br>LOW | CRITICAL       |

<sup>&</sup>lt;sup>7</sup> Considerable heterogeneity (I2>80%)

<sup>&</sup>lt;sup>8</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm <sup>9</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

| Quality              | assessment           |                                  |                             |                            |                              |                             | No of pat                      | ients  | Effect                      |  |             |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|--------------------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias                  | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Trauma<br>-<br>focuse<br>d CBT | Eye movement desensitisation and reprocessing (EMDR) | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Qualit<br>y | Importanc<br>e |
|                      |                      |                                  |                             |                            |                              |                             |                                |  |                             | (0.66<br>lower to<br>0.19<br>higher)                         |             |                |
| PTSD s               | symptomatolog        | gy cliniciar                     | n-rated (follow-u           | o mean 8 week              | s; measured v                | vith: CAPS-CA ch            | ange score                     | e; Better indicated                                  | by lower v                  | alues)   |             |                |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 23                             | 25   | -                           | SMD<br>0.04<br>higher<br>(0.53<br>lower to<br>0.6<br>higher) | VERY<br>LOW | CRITICAL       |
| Emotic               | nal and behav        | ioural prob                      | olems at endpoir            | it (follow-up m            | ean 6 weeks;                 | measured with: S            | DQ-A chan                      | ge score; Better in                                  | dicated by                  | / lower valu   | es)         |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup>     | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup>         | reporting bias <sup>3</sup> | 42                             | 43   | -                           | SMD<br>0.55<br>higher<br>(0.12 to<br>0.99<br>higher)         | VERY<br>LOW | IMPORTA<br>NT  |
| Emotio               | nal and behav        | ioural prob                      | olems at 3-month            | follow-up (fol             | low-up mean :                | 3 months; measu             | red with: S                    | DQ-A change scor                                     | e; Better iı                | ndicated by  | lower va    | lues)          |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup>     | no serious<br>inconsistency | no serious indirectness    | serious <sup>5</sup>         | reporting bias <sup>3</sup> | 42                             | 43   | -                           | SMD<br>0.46<br>higher<br>(0.03 to<br>0.89<br>higher)         | VERY<br>LOW | IMPORTA<br>NT  |

| Quality              | assessment           |                              |                             |                            |                      |                             | No of pat             | 1  | Effect                      |  |             |                |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Trauma - focuse d CBT | Eye movement desensitisation and reprocessing (EMDR) | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.45<br>higher<br>(0.02 to<br>0.89<br>higher)         | VERY<br>LOW | IMPORTA<br>NT  |
| Quality              | of life at endp      | oint (follow                 | v-up mean 6 wee             | eks; measured              | with: KIDSCR         | EEN-27: Global H            | RQoL T-so             | cores; change scor                                   | e; Better i                 | ndicated by  | higher v    | alues)         |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.23<br>lower<br>(0.66<br>lower to<br>0.2<br>higher)  | VERY<br>LOW | IMPORTA<br>NT  |
| •                    |                      | onth follow                  | -up (follow-up m            | ean 3 months;              | measured wit         | h: KIDSCREEN-2              | 7: Global F           | IRQoL T-scores; ch                                   | nange sco                   | re; Better in  | dicated I   | y higher       |
| values)              | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD<br>0.39<br>lower<br>(0.82<br>lower to<br>0.04<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Quality values)      |                      | onth follow                  | v-up (follow-up r           | mean 12 month              | s; measured v        | with: KIDSCREEN             | l-27: Globa           | I HRQoL T-scores;                                    | change s                    | core; Better   | indicate    | d by higher    |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>6</sup> | reporting bias <sup>3</sup> | 42                    | 43   | -                           | SMD 0.3<br>lower<br>(0.73                                    | VERY<br>LOW | IMPORTA<br>NT  |

| Quality        | assessment           |                                  |                             |                         |                              |                       | No of pat                      | tients   | Effect                      |  |             |                |
|----------------|----------------------|----------------------------------|-----------------------------|-------------------------|------------------------------|-----------------------|--------------------------------|--|-----------------------------|--|-------------|----------------|
| No of studi es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s        | Imprecisio<br>n              | Other consideration s | Trauma<br>-<br>focuse<br>d CBT | Eye movement desensitisation and reprocessing (EMDR) | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Qualit<br>y | Importanc<br>e |
|                |                      |                                  |                             |                         |                              |                       |                                |  |                             | lower to<br>0.12<br>higher)                                  |             |                |
| Discon         | tinuation (follo     | ow-up 6-8 w                      | veeks; assessed             | with: Number            | of participants              | s lost to follow-up   | o for any re                   | ason)  |                             |  |             |                |
| 2              | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious indirectness | very<br>serious <sup>4</sup> | none                  | 6/65<br>(9.2%)                 | 8/68<br>(11.8%)                                      | RR 0.8<br>(0.31 to<br>2.05) | 24 fewer<br>per 1000<br>(from 81<br>fewer to<br>124<br>more) | LOW         | CRITICAL       |

CAPS=Clinician Administered PTSD Symptom;; CBT=cognitive behavioural therapy; Cl=confidence interval; CRTI= Children's Response to Trauma Inventory; EMDR=Eye Movement Desensitisation and Reprocessing; HRQoL=Health-Related Quality of Life; KIDSCREEN-27= Health-related quality of life questionnaire for children, young people and their parents; PTSD=post-traumatic stress disorder; RR=risk ratio; SDQ-A= Strength and Difficulties Questionnaires; SMD=standard mean difference

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>5</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>6</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

Trauma-focused CBT versus combined somatic and cognitive therapies for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                              |                             |                            |                      |                       | No of pati                | ents                                     | Effect                      |   |             |            |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------|---------------------------|--|-----------------------------|---|-------------|------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias              | Inconsistency               | Indirectness               | Imprecisio<br>n      | Other consideration s | Trauma-<br>focused<br>CBT | Combined somatic and cognitive therapies | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importance |
| PTSD s               | ymptomatolog         | y self-rate                  | d at endpoint (fol          | low-up mean 2              | weeks; meas          | ured with: SPTSS      | change sc                 | ore; Better ind                          | icated by le                | ower values)                                      |             |            |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 19                        | 20                                       | -                           | SMD 0.87<br>higher<br>(0.21 to<br>1.53<br>higher) | VERY<br>LOW | CRITICAL   |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 3-month follo          | ow-up (follow-u            | p mean 3 mor         | nths; measured w      | ith: SPTSS                | change score;                            | <b>Better ind</b>           | icated by low                                     | er values   | s)         |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | none                  | 19                        | 20                                       | -                           | SMD 0.8<br>higher<br>(0.15 to<br>1.46<br>higher)  | VERY<br>LOW | CRITICAL   |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 6-month follo          | ow-up (follow-u            | p mean 6 mor         | nths; measured w      | ith: SPTSS                | change score;                            | <b>Better ind</b>           | icated by low                                     | er values   | s)         |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | none                  | 19                        | 20                                       | -                           | SMD 0.83<br>higher<br>(0.17 to<br>1.48<br>higher) | VERY<br>LOW | CRITICAL   |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 12-month fol           | low-up (follow-            | up mean 12 m         | onths; measured       | with: SPTS                | S change sco                             | re; Better i                | ndicated by le                                    | ower valu   | ies)       |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 19                        | 20                                       | -                           | SMD 0.92<br>higher<br>(0.26 to<br>1.58<br>higher) | VERY<br>LOW | CRITICAL   |

| Quality              | assessment           |                              |                             |                            |                              |                       | No of pati                | ients                                    | Effect                      |  |             |               |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------|---------------------------|--|-----------------------------|--|-------------|---------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias              | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other consideration s | Trauma-<br>focused<br>CBT | Combined somatic and cognitive therapies | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importance    |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>         | none                  | 19                        | 20                                       | -                           | SMD 1.01<br>higher<br>(0.34 to<br>1.68<br>higher)          | VERY<br>LOW | IMPORTA<br>NT |
| Anxiety              | symptoms at          | 3-month fo                   | ollow-up (follow-ເ          | ıp mean 3 mont             | ths; measured                | d with: HADS-A ch     | nange score               | e; Better indica                         | ited by low                 | er values)   |             |               |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>         | none                  | 19                        | 20                                       | -                           | SMD 0.91<br>higher<br>(0.25 to<br>1.57<br>higher)          | VERY<br>LOW | IMPORTA<br>NT |
| <b>Anxiety</b>       | symptoms at          | 6-month fo                   | ollow-up (follow-u          | ıp mean 6 mont             | ths; measured                | d with: HADS-A ch     | nange score               | e; Better indica                         | ited by low                 | er values)   |             |               |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | serious <sup>3</sup>         | none                  | 19                        | 20                                       | -                           | SMD 0.22<br>higher<br>(0.41<br>lower to<br>0.85<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| <b>Anxiety</b>       | symptoms at          | 12-month                     | follow-up (follow-          | -up mean 12 mo             | onths; measu                 | red with: HADS-A      | change sco                | ore; Better ind                          | icated by l                 | ower values)   |             |               |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious<br>indirectness | very<br>serious <sup>4</sup> | none                  | 19                        | 20                                       | -                           | SMD 0.09<br>lower<br>(0.71<br>lower to<br>0.54<br>higher)  | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment           |                              |                             |                            |                              |                       | No of pati                | ents                                     | Effect                      |  |             |                |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------|---------------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias                 | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other consideration s | Trauma-<br>focused<br>CBT | Combined somatic and cognitive therapies | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | none                  | 19                        | 20                                       | -                           | SMD 1.3<br>higher (0.6<br>to 1.99<br>higher)               | VERY<br>LOW | IMPORTA<br>NT  |
| •                    |                      |                              |                             |                            |                              | ured with: HADS-      |                           | -  | dicated by                  |  | )           |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>         | none                  | 19                        | 20                                       | -                           | SMD 0.45<br>higher<br>(0.19<br>lower to<br>1.09<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depress              | sion symptoms        | s at 6-mon                   | th follow-up (follo         | ow-up mean 6 n             | nonths; meas                 | ured with: HADS-      | D change s                | core; Better in                          | dicated by                  | lower values   | )           |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>3</sup>         | none                  | 19                        | 20                                       | -                           | SMD 0.3<br>higher<br>(0.33<br>lower to<br>0.93<br>higher)  | VERY<br>LOW | IMPORTA<br>NT  |
| Depress              | sion symptoms        | s at 12-moi                  | nth follow-up (fol          | low-up mean 12             | 2 months; me                 | asured with: HAD      | S-D change                | score; Better                            | indicated                   | by lower valu  | es)         |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | none                  | 19                        | 20                                       | -                           | SMD 0.66<br>higher<br>(0.02 to<br>1.31<br>higher)          | VERY<br>LOW | IMPORTA<br>NT  |
| Discont              | inuation (follo      | w-up mear                    | 2 weeks; assess             | sed with: Numb             | er of participa              | ants lost to follow   | -up for any               | reason)                                  |                             |  |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>         | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>4</sup> | none                  | 1/20<br>(5%)              | 0/20<br>(0%)                             | RR 3<br>(0.13 to<br>69.52)  | -  | VERY<br>LOW | CRITICAL       |

CBT=cognitive behavioural therapy; CI=confidence interval; HADS-A/D= Hospital Anxiety and Depression Scale-Anxiety/Depression; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; SPTSS= Screen for Post-Traumatic Stress Symptoms

Trauma-focused CBT + parent training versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                      |                             |                         |                      |                             | No of patie                                       | nts          | Effect                  |   |             |                |
|----------------------|----------------------|----------------------|-----------------------------|-------------------------|----------------------|-----------------------------|---|--------------|-------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency               | Indirectness            | Imprecisio<br>n      | Other considerations        | Trauma-<br>focused<br>CBT +<br>parent<br>training | Waitlis<br>t | Relative<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| PTSD s               | ymptomatolog         | y clinician-         | rated at endpoint           | (follow-up mea          | n 20 weeks; m        | neasured with: AD           | IS-C: PTSD;                                       | change so    | ore; Better             | r indicated by                                      | lower val   | lues)          |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 1.73<br>lower (2.69<br>to 0.77<br>lower)        | VERY<br>LOW | CRITICAL       |
| PTSD s values)       | ymptomatolog         | y clinician-         | rated at 3-month            | follow-up (follo        | w-up mean 3 r        | nonths; measured            | with: ADIS-                                       | C: PTSD;     | change sco              | ore; Better ind                                     | icated by   | lower          |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 1.34<br>lower (2.24<br>to 0.44<br>lower)        | VERY<br>LOW | CRITICAL       |
| Anxiety              | symptoms at          | endpoint (f          | ollow-up mean 20            | weeks; measu            | red with: RCM        | AS; change score            | ; Better indic                                    | cated by lo  | ower values             | s)  |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.33<br>lower (1.13<br>lower to<br>0.48 higher) | VERY<br>LOW | IMPORTA<br>NT  |

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

| Quality             | assessment           |                      |                             |                            |                      |                             | No of patie                                       | ents         | Effect                  |   |             |                |
|---------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|---|--------------|-------------------------|---|-------------|----------------|
| No of studie s      | Design               | Risk of bias         | Inconsistency               | Indirectness               | Imprecisio<br>n      | Other considerations        | Trauma-<br>focused<br>CBT +<br>parent<br>training | Waitlis<br>t | Relative<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| 1                   | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.75<br>lower (1.58<br>lower to<br>0.09 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depres              | sion symptom         | s at endpoir         | nt (follow-up mea           | n 20 weeks; me             | asured with: (       | CDI; change score           | ; Better indi                                     | cated by lo  | ower values             | s)  |             |                |
| 1                   | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.61<br>lower (1.43<br>lower to<br>0.21 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depress             | sion symptom         | s at 3-montl         | n follow-up (follow         | w-up mean 3 mo             | onths; measu         | red with: CDI; char         | nge score; B                                      | etter indic  | cated by lov            | wer values)   |             |                |
| 1                   | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious<br>indirectness | serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.36<br>lower (1.17<br>lower to<br>0.45 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Emotion lower value |                      | oural probl          | ems-Internalizing           | at endpoint (fo            | llow-up mean         | 20 weeks; measu             | red with: CB                                      | CL: Intern   | alizing; cha            | ange score; Be                                      | etter indi  | cated by       |
| 1                   | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.45<br>lower (1.26<br>lower to<br>0.36 higher) | VERY<br>LOW | IMPORTA<br>NT  |
|                     | nal and behavi       |                      | ems-Internalizing           | at 3-month follo           | ow-up (follow        | -up mean 3 month            | s; measured                                       | d with: CB   | CL: Interna             | lizing; change                                      | score; B    | etter          |
| 1                   | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 12  | 12           | -                       | SMD 0.92<br>lower (1.77                             | VERY<br>LOW | IMPORTA<br>NT  |

| Quality              | assessment                        |                                  |                             |                         |                              |                             | No of patie                                       | nts                 | Effect                     |   |             |                |
|----------------------|-----------------------------------|----------------------------------|-----------------------------|-------------------------|------------------------------|-----------------------------|---|---------------------|----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design                            | Risk of bias                     | Inconsistency               | Indirectness            | Imprecisio<br>n              | Other considerations        | Trauma-<br>focused<br>CBT +<br>parent<br>training | Waitlis<br>t        | Relative<br>(95%<br>CI)    | Absolute  | Qualit<br>y | Importanc<br>e |
| Emotio<br>lower v    |                                   | oural proble                     | ems-Externalizing           | g at endpoint (fo       | ollow-up mear                | n 20 weeks; meası           | ured with: CE                                     | CL: Exter           | nalizing; c                | hange score; I                                      | Better inc  | licated by     |
| 1                    | randomised<br>trials              | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness | serious <sup>4</sup>         | reporting bias <sup>3</sup> | 12  | 12                  | -                          | SMD 0.44<br>lower (1.25<br>lower to<br>0.37 higher) | VERY<br>LOW | IMPORTA<br>NT  |
|                      | nal and behavi<br>ed by lower val |                                  | ems-Externalizino           | g at 3-month fol        | low-up (follow               | /-up mean 3 mont            | hs; measure                                       | d with: CB          | CL: Extern                 | alizing; chang                                      | e score;    | Better         |
| 1                    | randomised<br>trials              | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12                  | -                          | SMD 0.88<br>lower (1.73<br>to 0.04<br>lower)        | VERY<br>LOW | IMPORTA<br>NT  |
| Global               | functioning at                    | endpoint (fo                     | ollow-up mean 20            | weeks; measu            | red with: GAF                | ; change score; B           | etter indicate                                    | ed by high          | er values)                 |   |             |                |
| 1                    | randomised<br>trials              | serious <sup>1</sup>             | no serious inconsistency    | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12                  | -                          | SMD 2.02<br>higher (1.01<br>to 3.04<br>higher)      | VERY<br>LOW | IMPORTA<br>NT  |
| Global               | functioning at                    | 3-month fol                      | low-up (follow-up           | mean 3 month            | s; measured v                | with: GAF; change           | e score; Bette                                    | er indicate         | ed by highe                | r values)   |             |                |
| 1                    | randomised<br>trials              | serious <sup>1</sup>             | no serious inconsistency    | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12                  | -                          | SMD 2.04<br>higher (1.02<br>to 3.06<br>higher)      | VERY<br>LOW | IMPORTA<br>NT  |
| Discon               | tinuation (follo                  | w-up mean                        | 20 weeks; assess            | sed with: Numb          | er of participa              | ints lost to follow-        | up for any re                                     | ason)               |                            |   |             |                |
| 1                    | randomised<br>trials              | no<br>serious<br>risk of<br>bias | no serious inconsistency    | no serious indirectness | very<br>serious <sup>5</sup> | none                        | 3/12<br>(25%)                                     | 2/12<br>(16.7%<br>) | RR 1.5<br>(0.3 to<br>7.43) | 83 more per<br>1000 (from<br>117 fewer              | LOW         | CRITICAL       |

| Quality        | assessment |              |               |              |                 |                      | No of patie                                       | nts          | Effect                  |                  |             |                |
|----------------|------------|--------------|---------------|--------------|-----------------|----------------------|---|--------------|-------------------------|------------------|-------------|----------------|
| No of studie s | Design     | Risk of bias | Inconsistency | Indirectness | Imprecisio<br>n | Other considerations | Trauma-<br>focused<br>CBT +<br>parent<br>training | Waitlis<br>t | Relative<br>(95%<br>CI) | Absolute         | Qualit<br>y | Importanc<br>e |
|                |            |              |               |              |                 |                      |   |              |                         | to 1000<br>more) |             |                |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBT=cognitive behavioural therapy; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS= Revised Children's Manifest Anxiety Scale; RR=risk ratio; SMD=standardised mean difference

# Trauma-focused CBT + parent training versus trauma-focused CBT (child only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

|                | assessment   |                      |                          |                         |                      |                             | No of patie                                       | nts  | Effect                      |  |             |                |
|----------------|--|----------------------|--------------------------|-------------------------|----------------------|-----------------------------|---|--|-----------------------------|--|-------------|----------------|
| No of studie s | Design   | Risk of bias         | Inconsistency            | Indirectness            | Imprecisio<br>n      | Other consideration s       | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| PTSD s         | PTSD symptomatology clinician-rated at endpoint (follow-up mean 20 weeks; measured v |                      |                          |                         |                      | measured with: Al           | DIS-C: PTSD;                                      | change sco                                   | ore; Better                 | indicated by   | lower val   | lues)          |
| 1              | randomised<br>trials   | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.36<br>lower (1.16<br>lower to<br>0.45<br>higher) | VERY<br>LOW | CRITICAL       |

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>5</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

| Quality              | assessment           |                      |                             |                         |                              |                             | No of patie                                       | ents   | Effect                      |   |             |               |
|----------------------|----------------------|----------------------|-----------------------------|-------------------------|------------------------------|-----------------------------|---|--|-----------------------------|---|-------------|---------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency               | Indirectness            | Imprecisio<br>n              | Other consideration s       | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importance    |
| PTSD s<br>values)    | ymptomatolog         | y clinician          | -rated at 3-month           | follow-up (follo        | w-up mean 3                  | months; measure             | ed with: ADIS                                     | -C: PTSD; c                                  | hange sco                   | re; Better ind  | cated by    | lower         |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.48<br>lower (1.29<br>lower to<br>0.34<br>higher)  | VERY<br>LOW | CRITICAL      |
| Anxiety              | symptoms at          | endpoint (f          | ollow-up mean 20            | ) weeks; measu          | red with: RCI                | MAS; change sco             | re; Better ind                                    | licated by lo                                | wer values                  | s)  |             |               |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.14<br>higher<br>(0.66 lower<br>to 0.94<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| Anxiety              | symptoms at          | 3-month fo           | llow-up (follow-u           | p mean 3 montl          | ns; measured                 | with: RCMAS; ch             | ange score;                                       | Better indica                                | ated by lov                 | ver values)   |             |               |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.03<br>higher<br>(0.77 lower<br>to 0.83<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| Depres               | sion symptom         | s at endpoi          | nt (follow-up mea           | n 20 weeks; me          | easured with:                | CDI; change scor            | e; Better ind                                     | icated by lo                                 | wer values                  | )   |             |               |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.29<br>lower (1.09<br>lower to<br>0.52<br>higher)  | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment                    |                      |                             |                            |                              |                             | No of patie                                       | ents   | Effect                      |  |             |               |
|----------------------|-------------------------------|----------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|---|--|-----------------------------|--|-------------|---------------|
| No of<br>studie<br>s | Design                        | Risk of bias         | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other consideration s       | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importance    |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.07<br>lower (0.87<br>lower to<br>0.73<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| Emotion lower v      |                               | oural probl          | ems-Internalizing           | at endpoint (fo            | ollow-up mear                | 20 weeks; measi             | ured with: Cl                                     | BCL: Interna                                 | lizing; cha                 | inge score; Be   | etter indi  | cated by      |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.29<br>lower (1.1<br>lower to<br>0.51<br>higher)  | VERY<br>LOW | IMPORTA<br>NT |
|                      | nal and behavied by lower val |                      | ems-Internalizing           | at 3-month fol             | low-up (follow               | v-up mean 3 mont            | hs; measure                                       | d with: CBC                                  | L: Internal                 | izing; change  | score; B    | etter         |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.15<br>lower (0.95<br>lower to<br>0.66<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| Emotion lower v      |                               | oural probl          | ems-Externalizin            | g at endpoint (f           | ollow-up mea                 | n 20 weeks; meas            | ured with: C                                      | BCL: Extern                                  | alizing; ch                 | nange score; E   | Better inc  | licated by    |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.79<br>lower (1.63<br>lower to<br>0.04<br>higher) | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment           |                                  |                             |                            |                              |                             | No of patie                                       | ents   | Effect                      |   |             |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|---|--|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias                     | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other consideration s       | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness    | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.14<br>lower (0.94<br>lower to<br>0.67<br>higher)    | VERY<br>LOW | IMPORTA<br>NT  |
| Global f             | functioning at       | endpoint (f                      | ollow-up mean 20            | ) weeks; measu             | red with: GAF                | ; change score; I           | Better indica                                     | ted by highe                                 | er values)                  |   |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.3<br>higher (0.5<br>lower to<br>1.11<br>higher)     | VERY<br>LOW | IMPORTA<br>NT  |
| Global               | functioning at       | 3-month fo                       | llow-up (follow-up          | o mean 3 month             | ns; measured                 | with: GAF; chang            | e score; Bet                                      | ter indicated                                | by higher                   | values)   |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 12  | 12   | -                           | SMD 0.66<br>higher<br>(0.16 lower<br>to 1.49<br>higher)   | VERY<br>LOW | IMPORTA<br>NT  |
| Discont              | inuation (follo      | w-up mean                        | 20 weeks; asses             | sed with: Numb             | er of particip               | ants lost to follow         | -up for any i                                     | reason)                                      |                             |   |             |                |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>4</sup> | none                        | 3/12<br>(25%)                                     | 3/12<br>(25%)                                | RR 1<br>(0.25 to<br>4)      | 0 fewer per<br>1000 (from<br>188 fewer<br>to 750<br>more) | LOW         | CRITICAL       |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS=; RR=risk ratio; SMD=standardised mean difference

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains <sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

#### Trauma-focused CBT versus parent training (CBT with parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                      |                          |                         |                      |                             | No of pati                | ents   | Effect                      |  |             |            |
|----------------------|----------------------|----------------------|--------------------------|-------------------------|----------------------|-----------------------------|---------------------------|--|-----------------------------|--|-------------|------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency            | Indirectness            | Imprecisio<br>n      | Other considerations        | Trauma-<br>focused<br>CBT | Parent<br>training<br>(CBT<br>with<br>parent-<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importance |
| PTSD s               | ymptomatolog         | y clinician          | -rated at endpoint       | t (follow-up mea        | an 12 weeks; n       | neasured with: K-           | SADS-E: PT                | SD; change   | score; Be                   | tter indicated                                       | by lower    | values)    |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.34<br>lower (0.96<br>lower to<br>0.27 higher)  | VERY<br>LOW | CRITICAL   |
| PTSD s values)       |                      | y clinician          | -rated at 3-month        | follow-up (follo        | w-up mean 3          | months; measured            | d with: K-SA              | ADS-E: PTSE  | ); change                   | score; Better i                                      | ndicated    | by lower   |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>4</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.12<br>higher (0.49<br>lower to<br>0.73 higher) | VERY<br>LOW | CRITICAL   |
|                      | • •                  | y clinician          | -rated at 6-month        | follow-up (follo        | w-up mean 6          | months; measure             | d with: K-SA              | ADS-E: PTSE  | ); change                   | score; Better i                                      | ndicated    | by lower   |
| values)              | randomised           | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.25<br>lower (0.87<br>lower to                  | VERY<br>LOW | CRITICAL   |

 <sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes
 <sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

| Quality              | assessment           |                      |                             |                            |                              |                             | No of pati                | ents   | Effect                      |  | _           |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|---------------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other considerations        | Trauma-<br>focused<br>CBT | Parent<br>training<br>(CBT<br>with<br>parent-<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.07<br>higher (0.54<br>lower to<br>0.68 higher) | VERY<br>LOW | CRITICAL       |
| PTSD syvalues)       | ymptomatolog         | y clinician-         | rated at 2-year fo          | llow-up (follow-           | up mean 2 ye                 | ars; measured wit           | h: K-SADS-                | E: PTSD; ch  | ange scoi                   | re; Better indic                                     | ated by I   | ower           |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup>         | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.64<br>higher (0.01<br>to 1.27<br>higher)       | VERY<br>LOW | CRITICAL       |
| Emotion lower va     |                      | oural probl          | lems-Externalizin           | g at endpoint (f           | ollow-up mea                 | n 12 weeks; meas            | ured with: C              | BCL Extern   | alizing ch                  | ange score; Be                                       | etter indi  | cated by       |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 20                        | 18   | -                           | SMD 0.13<br>higher (0.51<br>lower to<br>0.77 higher) | VERY<br>LOW | IMPORTA<br>NT  |
|                      | nal and behavi       |                      | lems-Externalizin           | g at 3-month fo            | llow-up (follow              | v-up mean 3 mont            | hs; measur                | ed with: CB  | CL Externa                  | alizing change                                       | score; B    | etter          |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 20                        | 18   | -                           | SMD 0.61<br>lower (1.27<br>lower to<br>0.04 higher)  | VERY<br>LOW | IMPORTA<br>NT  |
|                      | nal and behavi       |                      | lems-Externalizin           | g at 6-month fo            | llow-up (follow              | v-up mean 6 mont            | hs; measur                | ed with: CB  | CL Externa                  | alizing change                                       | score; B    | etter          |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>6</sup>         | reporting bias <sup>3</sup> | 20                        | 18   | -                           | SMD 0.75<br>lower (1.41                              | VERY<br>LOW | IMPORTA<br>NT  |

| Quality              | assessment                        |                      |                             |                            |                      |                             | No of pati                | ents   | Effect                      |   |             |                |
|----------------------|-----------------------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|---------------------------|--|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design                            | Risk of bias         | Inconsistency               | Indirectness               | Imprecisio<br>n      | Other considerations        | Trauma-<br>focused<br>CBT | Parent<br>training<br>(CBT<br>with<br>parent-<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
|                      |                                   |                      |                             |                            |                      |                             |                           |  |                             | to 0.09<br>lower)                                   |             |                |
|                      | nal and behavi<br>ed by lower val |                      | lems-Externalizin           | g at 12-month f            | ollow-up (follo      | ow-up mean 12 mo            | nths; meas                | ured with: (   | CBCL Exte                   | rnalizing chan                                      | ge score    | ; Better       |
| 1                    | randomised<br>trials              | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup> | reporting bias <sup>3</sup> | 20                        | 18   | -                           | SMD 0.79<br>lower (1.45<br>to 0.12<br>lower)        | VERY<br>LOW | IMPORTA<br>NT  |
|                      | nal and behavi<br>er values)      | oural prob           | lems-Externalizin           | g at 2-year follo          | w-up (follow-        | up mean 2 years; ı          | measured w                | vith: CBCL E   | Externalizi                 | ng change sco                                       | re; Bette   | r indicated    |
| 1                    | randomised<br>trials              | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20                        | 18   | -                           | SMD 0.53<br>lower (1.18<br>lower to<br>0.12 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depres               | sion symptoms                     | s at endpoi          | nt (follow-up mea           | an 12 weeks; me            | easured with:        | CDI change score            | ; Better indi             | cated by lo  | wer values                  | s)  |             |                |
| 1                    | randomised<br>trials              | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.3<br>higher (0.32<br>lower to<br>0.92 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depres               | sion symptom                      | s at 3-mont          | th follow-up (follo         | w-up mean 3 m              | onths; measu         | red with: CDI chai          | nge score; E              |  | ated by low                 | ver values)   |             |                |
| 1                    | randomised<br>trials              | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.12<br>higher (0.49<br>lower to                | VERY<br>LOW | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                         |                              |                             | No of pati                | ents   | Effect                      |  |             |                |
|----------------------|----------------------|----------------------|-----------------------------|-------------------------|------------------------------|-----------------------------|---------------------------|--|-----------------------------|--|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency               | Indirectness            | Imprecisio<br>n              | Other considerations        | Trauma-<br>focused<br>CBT | Parent<br>training<br>(CBT<br>with<br>parent-<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.09<br>higher (0.53<br>lower to 0.7<br>higher)  | VERY<br>LOW | IMPORTA<br>NT  |
| Depres               | sion symptoms        | at 12-moi            | nth follow-up (foll         | ow-up mean 12           | months; mea                  | sured with: CDI cl          | nange score               | ; Better ind   | icated by l                 | ower values)   |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>4</sup>         | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.31<br>higher (0.31<br>lower to<br>0.93 higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depres               | sion symptoms        | s at 2-year          | follow-up (follow-          | up mean 2 year          | s; measured                  | with: CDI change            | score; Bette              | r indicated  | by lower v                  | alues)   |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>6</sup>         | reporting bias <sup>3</sup> | 21                        | 20   | -                           | SMD 0.73<br>higher (0.1<br>to 1.37<br>higher)        | VERY<br>LOW | IMPORTA<br>NT  |

CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI=Children's Depression Inventory; CI=confidence interval; K-SADS-E=Kiddie Schedule for Affective Disorders and Schizophrenia-Epidemiological; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standard mean difference

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple outcomes

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

<sup>&</sup>lt;sup>5</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>6</sup> OIS not met (N<400)

Trauma-focused CBT (+ psychoeducational group) versus psychoeducational group for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality                 | y assessmen           | t                        |                                 |                                |                      |                      | No of patients                                  |                             | Effect                         |  |               |               |
|-------------------------|-----------------------|--------------------------|---------------------------------|--------------------------------|----------------------|----------------------|---|-----------------------------|--------------------------------|--|---------------|---------------|
| No<br>of<br>studi<br>es | Design                | Risk<br>of<br>bias       | Inconsisten<br>cy               | Indirectne<br>ss               | Imprecisi<br>on      | Other considerations | Trauma-focused CBT (+ psychoeducatio nal group) | Psychoeducatio<br>nal group | Relati<br>ve<br>(95%<br>CI)    | Absol<br>ute                                       | Quality       | Importa<br>ce |
| PTSD                    | symptomato            | logy self-               | rated at endpo                  | int (follow-up                 | mean 20 w            | eeks; measured       | with: UCLA PTSD-                                | RI; change score; B         | etter indi                     | cated by I   | ower values)  |               |
| 1                       | randomise<br>d trials | seriou<br>s <sup>1</sup> | no serious<br>inconsistenc<br>y | no serious<br>indirectnes<br>s | serious <sup>2</sup> | none                 | 66  | 61                          | -                              | SMD<br>0.46<br>lower<br>(0.81<br>to 0.11<br>lower) | LOW           | CRITICA<br>L  |
| PTSD                    | symptomato            | logy self-               | rated at 4 mon                  | th follow-up                   | (follow-up m         | ean 4 months;        | measured with: UCL                              | _A PTSD-RI; change          | score; B                       | etter indi   | cated by lowe | r values)     |
| 1                       | randomise<br>d trials | seriou<br>s <sup>1</sup> | no serious<br>inconsistenc<br>y | no serious<br>indirectnes<br>s | serious <sup>2</sup> | none                 | 36  | 29                          | -                              | SMD<br>0.57<br>lower<br>(1.07<br>to 0.07<br>lower) | LOW           | CRITICA<br>L  |
| Respo                   | onse at endpo         | oint (follo              | w-up mean 20                    | weeks; asses                   | ssed with: N         | umber of people      | e showing clinically                            | significant improve         | ement, ba                      | sed on re  | liable change | indices       |
| 1                       | randomise<br>d trials | seriou<br>s <sup>1</sup> | no serious<br>inconsistenc<br>y | no serious<br>indirectnes<br>s | serious <sup>3</sup> | none                 | 38/77<br>(49.4%)                                | 20/82<br>(24.4%)            | RR<br>2.02<br>(1.3 to<br>3.15) | more per 1000 (from 73 more to 524 more)           | LOW           | CRITICA<br>L  |

| Quality                 | assessmen             | t                                 |                                 |                                |                                     |                      | No of patients                                  |                            | Effect                             |  |              |                |
|-------------------------|-----------------------|-----------------------------------|---------------------------------|--------------------------------|-------------------------------------|----------------------|---|----------------------------|------------------------------------|--|--------------|----------------|
| No<br>of<br>studi<br>es | Design                | Risk<br>of<br>bias                | Inconsisten<br>cy               | Indirectne<br>ss               | Imprecisi<br>on                     | Other considerations | Trauma-focused CBT (+ psychoeducatio nal group) | Psychoeducatio nal group   | Relati<br>ve<br>(95%<br>CI)        | Absol<br>ute   | Quality      | Importan<br>ce |
| 1                       | randomise<br>d trials | seriou<br>s <sup>1</sup>          | no serious<br>inconsistenc<br>y | no serious<br>indirectnes<br>s | serious <sup>3</sup>                | none                 | 29/77<br>(37.7%)                                | 14/82<br>(17.1%)           | RR<br>2.21<br>(1.26<br>to<br>3.85) | 207<br>more<br>per<br>1000<br>(from<br>44<br>more<br>to 487<br>more) | LOW          | CRITICA<br>L   |
| Depres                  | randomise             | ms at en<br>seriou                | no serious                      | no serious                     | weeks; meas<br>serious <sup>2</sup> | none                 | RS change score; Be                             | etter indicated by love 60 | ver value                          | s)<br>SMD  |              | IMPORT         |
| l                       | d trials              | s <sup>1</sup>                    | inconsistenc<br>y               | indirectnes<br>s               | Serious                             | none                 | 00  | 00                         | -                                  | 0.44<br>lower<br>(0.8 to<br>0.09<br>lower)                           | LOW          | ANT            |
| Depres                  | sion sympto           | ms at 4 r                         | month follow-u                  | ıp (follow-up                  | mean 4 mon                          | iths; measured       | with: DSRS change                               | score; Better indica       | ted by lo                          | wer value  | s)           |                |
| 1                       | randomise<br>d trials | seriou<br>s <sup>1</sup>          | no serious<br>inconsistenc<br>y | no serious<br>indirectnes<br>s | serious <sup>2</sup>                | none                 | 36  | 30                         | -                                  | SMD<br>0.59<br>lower<br>(1.08<br>to 0.09<br>lower)                   | LOW          | IMPORT<br>ANT  |
| Discon                  | tinuation (fo         | llow-up r                         | nean 20 weeks                   | ; assessed w                   | vith: Number                        | r of participants    | lost to follow-up fo                            | r any reason)              |                                    |  |              |                |
| 1                       | randomise<br>d trials | no<br>seriou<br>s risk<br>of bias | no serious<br>inconsistenc<br>y | no serious indirectnes s       | serious <sup>4</sup>                | none                 | 11/77<br>(14.3%)                                | 21/82<br>(25.6%)           | RR<br>0.56<br>(0.29<br>to<br>1.08) | fewer<br>per<br>1000<br>(from  | MODERAT<br>E | CRITICA<br>L   |

| Quality                 | / assessmen | ıt                 |                   |                  |                 |                      | No of patients                                  |                          | Effect                      |                                |         |                |
|-------------------------|-------------|--------------------|-------------------|------------------|-----------------|----------------------|---|--------------------------|-----------------------------|--------------------------------|---------|----------------|
| No<br>of<br>studi<br>es | Design      | Risk<br>of<br>bias | Inconsisten<br>cy | Indirectne<br>ss | Imprecisi<br>on | Other considerations | Trauma-focused CBT (+ psychoeducatio nal group) | Psychoeducatio nal group | Relati<br>ve<br>(95%<br>CI) | Absol<br>ute                   | Quality | Importan<br>ce |
|                         |             |                    |                   |                  |                 |                      |   |                          |                             | 182<br>fewer<br>to 20<br>more) |         |                |

CBT=cognitive behavioural therapy; CI=confidence interval; DSRS= Depression Self-Rating Scale; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standard mean difference; UCLA PTSD-RI=UCLA PTSD-Reaction Index

#### Psychological: Non-trauma-focused CBT

Non-trauma focused CBT (+ TAU) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality        | assessment                       |                 |                          |                         |                 |                      | No of patie                                | ents     | Effect               |                         |             |                |
|----------------|----------------------------------|-----------------|--------------------------|-------------------------|-----------------|----------------------|--|----------|----------------------|-------------------------|-------------|----------------|
| No of studie s | Design                           | Risk of<br>bias | Inconsistency            | Indirectness            | Imprecisio<br>n | Other considerations | Non-<br>trauma<br>focused<br>CBT<br>(+TAU) | TAU      | Relative<br>(95% CI) | Absolute                | Qualit<br>v | Importanc<br>e |
|                | sion symptoms<br>ed by lower val |                 | t (follow-up mean        | 13 weeks; mea           | sured with: A   | dolescent Psycho     | pathology S                                | cale: Ax | kis I - Major        | Depression; cl          | nange sc    | ore; Better    |
| 1              | randomised<br>trials             | serious1        | no serious inconsistency | no serious indirectness | serious2        | reporting bias3      | 18   | 15       | -                    | SMD 0.33<br>lower (1.02 | VERY<br>LOW | IMPORTA<br>NT  |

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple outcomes

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> OIS not met (events<300)

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

| Quality              | assessment                       |                                  |  |                            |                  |                      | No of patie                                | ents              | Effect                       |  |             |               |
|----------------------|----------------------------------|----------------------------------|--|----------------------------|------------------|----------------------|--|-------------------|------------------------------|--|-------------|---------------|
| No of<br>studie<br>s | Design                           | Risk of<br>bias                  | Inconsistency                          | Indirectness               | Imprecisio<br>n  | Other considerations | Non-<br>trauma<br>focused<br>CBT<br>(+TAU) | TAU               | Relative<br>(95% CI)         | Absolute   | Qualit<br>y | Importance    |
|                      |                                  |                                  |  |                            |                  |                      |  |                   |                              | lower to 0.37 higher)                                  |             |               |
|                      | sion symptoms<br>Better indicate |                                  | h follow-up (follow<br>values)         | <i>ı</i> -up mean 3 mo     | nths; measure    | ed with: Adolescer   | nt Psychopa                                | thology           | Scale: Axis                  | s I - Major Depr                                       | ession; c   | hange         |
| 1                    | randomised<br>trials             | serious1                         | no serious<br>inconsistency            | no serious indirectness    | serious4         | reporting bias3      | 18   | 15                | -                            | SMD 0.71<br>higher (0 to<br>1.42 higher)               | VERY<br>LOW | IMPORTA<br>NT |
|                      |                                  |                                  | ns at endpoint (fol<br>y lower values) | low-up mean 13             | weeks; meas      | ured with: Adoles    | cent Psycho                                | patholo           | gy Scale: A                  | xis I - Substan  | ce Use D    | isorder;      |
| 1                    | randomised<br>trials             | serious1                         | no serious<br>inconsistency            | no serious<br>indirectness | serious5         | reporting bias3      | 18   | 15                | -                            | SMD 1.03<br>lower (1.77<br>to 0.3 lower)               | VERY<br>LOW | IMPORTA<br>NT |
|                      |                                  |                                  | ns at 3-month follo                    |                            | mean 3 mon       | ths; measured wit    | h: Adolesce                                | nt Psyc           | hopatholog                   | y Scale: Axis I  | - Substai   | nce Use       |
| 1                    | randomised<br>trials             | serious1                         | no serious<br>inconsistency            | no serious indirectness    | serious4         | reporting bias3      | 18   | 15                | -                            | SMD 0.63<br>higher (0.08<br>lower to 1.33<br>higher)   | VERY<br>LOW | IMPORTA<br>NT |
| Discont              | tinuation (follo                 | w-up mean                        | 13 weeks; assess                       | ed with: Numbe             | r of participan  | ts lost to follow-u  |  |                   |                              |  |             |               |
| 1                    | randomised<br>trials             | no<br>serious<br>risk of<br>bias | no serious inconsistency               | no serious indirectness    | very<br>serious6 | none                 | 4/18<br>(22.2%)                            | 3/15<br>(20<br>%) | RR 1.11<br>(0.29 to<br>4.21) | 22 more per<br>1000 (from<br>142 fewer to<br>642 more) | LOW         | CRITICAL      |

CBT=cognitive behavioural therapy; Cl=confidence interval; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; TAU=treatment as usual <sup>1</sup> Risk of bias is high or unclear across multiple outcomes

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

<sup>&</sup>lt;sup>5</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>6</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

## **Psychological: Psychodynamic therapies**

Child-parent psychotherapy using play versus parent training (case management and individual treatment for parent-only) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |              |                             |                            |                      |                             | No of patients                                  | i   | Effect                      |  |             |               |
|----------------------|----------------------|--------------|-----------------------------|----------------------------|----------------------|-----------------------------|---|---|-----------------------------|--|-------------|---------------|
| No of<br>studi<br>es | Design               | Risk of bias | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Child-parent<br>psychothera<br>py using<br>play | Parent<br>training (case<br>management<br>and individual<br>treatment for<br>parent-only) | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e                                       | Quali<br>ty | Importance    |
| PTSD s               | symptomatolog        | gy clinicia  | n-rated (follow-น           | p mean 50 we               | eks; measure         | d with: DC 0-3; cl          | nange score; Be                                 | etter indicated by  | lower valu                  | ıes)   |             |               |
| 1                    | randomised<br>trials | Serious<br>1 | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 36  | 29  | -                           | SMD<br>1.19<br>lower<br>(1.72 to<br>0.66<br>lower) | VERY<br>LOW | CRITICAL      |
| Emotio               | nal and behav        | ioural pro   | blems at endpoi             | nt (follow-up n            | nean 50 weeks        | s; measured with            | : CBCL total; ch                                | ange score; Bett  | er indicate                 | ed by lowe   | r values)   |               |
| 1                    | randomised<br>trials | serious<br>1 | no serious inconsistency    | no serious<br>indirectness | serious              | reporting bias <sup>3</sup> | 36  | 29  | -                           | SMD<br>0.79<br>lower<br>(1.3 to<br>0.28<br>lower)  | VERY<br>LOW | IMPORTA<br>NT |
| Emotic               | nal and behav        | ioural pro   | blems at 6-mont             | h follow-up (fo            | llow-up mean         | 6 months; meas              | ured with: CBC                                  | L total; change so  | ore; Bette                  | er indicated                                       | d by low    | er values)    |
| 1                    | randomised<br>trials | serious<br>1 | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 27  | 23  | -                           | SMD<br>0.98<br>lower<br>(1.58 to<br>0.39<br>lower) | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment           |                                  |                             |                            |                              |                       | No of patients                                  | <b>;</b>  | Effect                             |  |             |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------|---|---|------------------------------------|--|-------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s | Child-parent<br>psychothera<br>py using<br>play | Parent<br>training (case<br>management<br>and individual<br>treatment for<br>parent-only) | Relativ<br>e<br>(95%<br>CI)        | Absolut<br>e   | Quali<br>ty | Importanc<br>e |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>4</sup> | none                  | 6/42<br>(14.3%)                                 | 4/33<br>(12.1%)   | RR<br>1.18<br>(0.36<br>to<br>3.84) | 22 more<br>per<br>1000<br>(from 78<br>fewer to<br>344<br>more) | LOW         | CRITICAL       |

CBCL=Children's Behavioural Checklist; CI=confidence interval; DC=Diagnostic Criteria; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference;

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains <sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes <sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

## Psychological: Eye movement desensitisation and reprocessing (EMDR)

EMDR versus waitlist or TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                              |                             |                            |                              |                             | No of patients   |                        | Effect                      |   |             |            |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|--|------------------------|-----------------------------|---|-------------|------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias              | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Eye movement<br>desensitisation<br>and<br>reprocessing<br>(EMDR) | Waitlis<br>t or<br>TAU | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importance |
| PTSD s               | ymptomatolog         | gy self-rate                 | d at endpoint (fo           | llow-up 3-6 we             | eks; measure                 | d with: CRTI/CRIE           | S change score; B  | etter indic            | cated by lo                 | wer values)   |             |            |
| 2                    | randomised<br>trials | very<br>serious <sup>1</sup> | very serious2               | no serious<br>indirectness | very<br>serious <sup>3</sup> | reporting bias <sup>4</sup> | 53   | 29                     | -                           | SMD 0.9<br>lower<br>(2.64<br>lower to<br>0.85<br>higher)      | VERY<br>LOW | CRITICAL   |
| PTSD s               | symptomatolog        | gy self-rate                 | d at 2-month foll           | ow-up (follow-             | up mean 2 mo                 | nths; measured v            | with: CRIES change   | e score; B             | etter indic                 | ated by lowe  | er values   | )          |
|                      | randomised<br>trials | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup>         | none                        | 12   | 11                     | -                           | SMD<br>0.72<br>lower<br>(1.57<br>lower to<br>0.13<br>higher)  | LOW         | CRITICAL   |
| PTSD s               | symptomatolog        | gy clinician                 | -rated (follow-up           | mean 8 weeks               | s; measured w                | vith: PTSS-C char           | ige score; Better in   | dicated by             | y lower va                  | lues)   |             |            |
| 1                    | randomised<br>trials | serious <sup>1</sup>         | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>3</sup> | none                        | 17   | 16                     | -                           | SMD<br>0.07<br>higher<br>(0.61<br>lower to<br>0.76<br>higher) | VERY<br>LOW | CRITICAL   |

| Quanty               | / assessment         |                                  |                             |                            |                              |                             | No of patients                                       |                        | Effect                          |  |             |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|--|------------------------|---------------------------------|--|-------------|----------------|
| No of<br>studi<br>es | Design               | Risk of<br>bias                  | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Eye movement desensitisation and reprocessing (EMDR) | Waitlis<br>t or<br>TAU | Relativ<br>e<br>(95%<br>CI)     | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup>     | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup>         | reporting bias <sup>4</sup> | 43   | 18                     | -                               | SMD<br>1.52<br>lower<br>(2.14 to<br>0.91<br>lower)   | VERY<br>LOW | IMPORTA<br>NT  |
| Quality              | of life (follow-     | -up mean 6                       | weeks; measure              | ed with: KIDSC             | REEN-27: Glo                 | bal HRQoL T-sco             | res; change score;                                   | Better ind             | licated by                      | higher value   | es)         |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup>     | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>6</sup>         | reporting bias <sup>4</sup> | 43   | 18                     | -                               | SMD<br>0.81<br>higher<br>(0.24 to<br>1.38<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Discon               | tinuation (follo     | ow-up 3-8 v                      | veeks; assessed             | with: Number               | of participants              | s lost to follow-up         | for any reason)                                      |                        |                                 |  |             |                |
| 3                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>3</sup> | none                        | 5/74<br>(6.8%)                                       | 6/49<br>(12.2%<br>)    | RR<br>0.65<br>(0.15 to<br>2.88) | 43 fewer<br>per 1000<br>(from 104<br>fewer to<br>230 | LOW         | CRITICAL       |

CI=confidence interval; CRIES= Children's Revised Impact of Event Scale; CRTI= Children's Response to Trauma Inventory; HRQoL=Health Related Quality of Life; KIDSCREEN-27= Health-related quality of life questionnaire for children, young people and their parents; PTSD=post-traumatic stress disorder; PTSS=Post-Traumatic Stress Symptom; RR=risk ratio; SDQ-A= Strength and Difficulties Questionnaires; SMD=standardised mean difference; TAU=treatment as usual;

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> Considerable heterogeneity (I2>80%)

<sup>&</sup>lt;sup>3</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>4</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>5</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>6</sup> OIS not met (N<400)

## **Psychological: Combined somatic and cognitive therapies**

Combined somatic and cognitive therapies versus no treatment for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality assessment   |                      |                              |                             |                            |                      |                       | No of patients                           |                     | Effect                      |   |             |                |
|----------------------|----------------------|------------------------------|-----------------------------|----------------------------|----------------------|-----------------------|--|---------------------|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias              | Inconsistency               | Indirectness               | Imprecisio<br>n      | Other consideration s | Combined somatic and cognitive therapies | No<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| PTSD s               | ymptomatolog         | y self-rate                  | d at endpoint (fol          | low-up mean 2              | weeks; measi         | ured with: SPTSS      | change score;                            | Better indic        | cated by lo                 | wer values)                                     |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.85<br>lower (2.6<br>to 1.1<br>lower)      | VERY<br>LOW | CRITICAL       |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 3-month follo          | w-up (follow-up            | p mean 3 mon         | ths; measured w       | th: SPTSS cha                            | nge score; E        | Better indi                 | cated by low                                    | er values   | s)             |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.96<br>lower<br>(2.72 to<br>1.19<br>lower) | VERY<br>LOW | CRITICAL       |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 6-month follo          | w-up (follow-up            | p mean 6 mon         | ths; measured wi      | th: SPTSS cha                            | nge score; E        | Better indi                 | cated by low                                    | er values   | 5)             |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.3<br>lower<br>(1.99 to<br>0.61<br>lower)  | VERY<br>LOW | CRITICAL       |
| PTSD s               | ymptomatolog         | y self-rate                  | d at 12-month fol           | low-up (follow-u           | up mean 12 m         | onths; measured       | with: SPTSS c                            | hange score         | e; Better ir                | ndicated by le                                  | ower valu   | ies)           |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.85<br>lower (2.6<br>to 1.1<br>lower)      | VERY<br>LOW | CRITICAL       |

| Quality              | assessment           |                              |                             |                         |                      |                       | No of patient                            | s                   | Effect                      |   |             |                |
|----------------------|----------------------|------------------------------|-----------------------------|-------------------------|----------------------|-----------------------|--|---------------------|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias              | Inconsistency               | Indirectness            | Imprecisio<br>n      | Other consideration s | Combined somatic and cognitive therapies | No<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| Anxiety              | symptoms at          | endpoint (                   | follow-up mean 2            | weeks; measu            | red with: HAD        | S-A change score      | e; Better indica                         | ted by lowe         | r values)                   |   |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 0.95<br>lower<br>(1.61 to<br>0.3 lower)     | VERY<br>LOW | IMPORTA<br>NT  |
| <b>Anxiety</b>       | symptoms at          | 3-month fo                   | ollow-up (follow-u          | ıp mean 3 mont          | ths; measured        | l with: HADS-A ch     | ange score; B                            | etter indicat       | ed by low                   | er values)                                      |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 0.89<br>lower<br>(1.54 to<br>0.24<br>lower) | VERY<br>LOW | IMPORTA<br>NT  |
| Anxiety              | symptoms at          | 6-month fo                   | llow-up (follow-u           | ıp mean 6 mont          | ths; measured        | with: HADS-A ch       | ange score; B                            | etter indicat       | ed by low                   | er values)                                      |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.15<br>lower<br>(1.82 to<br>0.47<br>lower) | VERY<br>LOW | IMPORTA<br>NT  |
| <b>Anxiety</b>       | symptoms at          | 12-month f                   | follow-up (follow-          | up mean 12 mo           | onths; measur        | red with: HADS-A      | change score;                            | Better indic        | ated by lo                  | wer values)                                     |             |                |
| 1                    | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.19<br>lower<br>(1.86 to<br>0.51<br>lower) | VERY<br>LOW | IMPORTA<br>NT  |
| Depres               | sion symptom         | s at endpoi                  | int (follow-up me           | an 2 weeks; me          | asured with: I       | HADS-D change s       | core; Better in                          | dicated by le       | ower value                  | es)   |             |                |
| 1                    | randomised trials    | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 0.94<br>lower<br>(1.59 to                   | VERY<br>LOW | IMPORTA<br>NT  |

| Quality        | assessment           |                              |                             |                         |                      |                       | No of patient                            | :s                  | Effect                      |   |             |                |
|----------------|----------------------|------------------------------|-----------------------------|-------------------------|----------------------|-----------------------|--|---------------------|-----------------------------|---|-------------|----------------|
| No of studie s | Design               | Risk of bias                 | Inconsistency               | Indirectness            | Imprecisio<br>n      | Other consideration s | Combined somatic and cognitive therapies | No<br>treatmen<br>t | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
|                |                      |                              |                             |                         |                      |                       |  |                     |                             | 0.28<br>lower)                                  |             |                |
| Depres         | sion symptoms        | s at 3-mon                   | th follow-up (follo         | ow-up mean 3 n          | nonths; meas         | ured with: HADS-      | D change scor                            | e; Better ind       | icated by                   | lower values                                    | )           |                |
| 1              | randomised trials    | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 0.75<br>lower (1.4<br>to 0.11<br>lower)     | VERY<br>LOW | IMPORTA<br>NT  |
| Depres         | sion symptoms        | at 6-mon                     | th follow-up (follo         | ow-up mean 6 n          | nonths; meas         | ured with: HADS-      | D change scor                            | e; Better ind       | icated by                   | lower values                                    | )           |                |
| 1              | randomised trials    | very<br>serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 0.85<br>lower (1.5<br>to 0.2<br>lower)      | VERY<br>LOW | IMPORTA<br>NT  |
| Depres         | sion symptoms        | at 12-moi                    | nth follow-up (fol          | low-up mean 12          | 2 months; mea        | asured with: HAD      | S-D change sc                            | ore; Better i       | ndicated k                  | y lower valu                                    | es)         |                |
| 1              | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency    | no serious indirectness | serious <sup>2</sup> | none                  | 20                                       | 20                  | -                           | SMD 1.38<br>lower<br>(2.07 to<br>0.68<br>lower) | VERY<br>LOW | IMPORTA<br>NT  |
| Discont        | tinuation (follo     | w-up mear                    | n 2 weeks; assess           | sed with: Numb          | er of participa      | ints lost to follow   | -up for any rea                          | son)                |                             |   |             |                |
| 1              | randomised           | serious1                     | no serious                  | no serious              | serious <sup>3</sup> | none                  | 0/20                                     | 0/20                | not                         | not pooled                                      |             | CRITICAL       |

CI=confidence interval; HADS-A/D= Hospital Anxiety and Depression Scale-Anxiety/Depression; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; SPTSS= Screen for Post-Traumatic Stress Symptoms; TAU=treatment as usual

1 Risk of bias is high or unclear across mutliple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400) <sup>3</sup> OIS not met (events<300)

### **Psychological: Supportive counselling**

Supportive counselling versus no treatment or waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of patien                      | its                                | Effect                      |  |                 |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------------------|------------------------------------|-----------------------------|--|-----------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Supportiv<br>e<br>counsellin<br>g | No<br>treatme<br>nt or<br>waitlist | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Quality         | Importane<br>e |
| PTSD s               | ymptomatolog         | gy self-rate                     | ed at endpoint (f           | ollow-up mean              | 6 weeks; me          | asured with: CRIE           | ES change sc                      | ore; Better i                      | ndicated b                  | oy lower va  | lues)           |                |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | none                        | 10                                | 12                                 | -                           | SMD<br>0.48<br>lower<br>(1.33<br>lower to<br>0.37<br>higher) | LOW             | CRITICAL       |
| PTSD s               | ymptomatolog         | gy self-rate                     | ed at 3-month fo            | llow-up (follow            | v-up mean 3 m        | onths; measured             | with: CRIES                       | change sco                         | re; Better                  | indicated b  | y lower values  | s)             |
| 1                    | randomised<br>trials | serious <sup>1</sup>             | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                        | 10                                | 12                                 | -                           | SMD<br>0.42<br>lower<br>(1.27<br>lower to<br>0.43<br>higher) | LOW             | CRITICAL       |
| PTSD s               | ymptomatolog         | gy clinicia                      | n-rated at 3-mon            | th follow-up (fe           | ollow-up mea         | n 3 months; meas            | sured with: CA                    | APS change                         | score; Be                   | etter indica   | ted by lower va | ılues)         |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 24                                | 28                                 | -                           | SMD<br>0.43<br>lower<br>(0.98<br>lower to<br>0.12<br>higher) | LOW             | CRITICAL       |

| Quality     | assessment           |                                  |                             |                            |                              |                             | No of patien                      | ıts                                | Effect                          |   |                |                |
|-------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|-----------------------------------|------------------------------------|---------------------------------|---|----------------|----------------|
| No of studi | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n              | Other consideration s       | Supportiv<br>e<br>counsellin<br>g | No<br>treatme<br>nt or<br>waitlist | Relativ<br>e<br>(95%<br>CI)     | Absolut<br>e  | Quality        | Importanc<br>e |
| 1           | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 23                                | 28                                 | -                               | SMD<br>0.11<br>lower<br>(0.66<br>lower to<br>0.44<br>higher)        | LOW            | CRITICAL       |
| PTSD s      | ymptomatolog         | gy clinicia                      | n-rated at 12-mo            | nth follow-up (            | follow-up mea                | an 12 months; me            | easured with:                     |                                    | ge score;                       | Better indi   | cated by lower | values)        |
| 1           | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>4</sup> | reporting bias <sup>3</sup> | 23                                | 28                                 | -                               | SMD 0<br>higher<br>(0.55<br>lower to<br>0.55<br>higher)             | VERY LOW       | CRITICAL       |
| Remiss      | ion at 12-mon        | th follow-ւ                      | up (follow-up me            | an 12 months;              | assessed wit                 | h: Number of peo            | ple no longei                     | r meeting di                       | agnostic (                      | criteria for  | PTSD)          |                |
| 1           | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4             | reporting bias <sup>3</sup> | 13/28<br>(46.4%)                  | 15/28<br>(53.6%)                   | RR<br>0.87<br>(0.51 to<br>1.47) | 70 fewer<br>per<br>1000<br>(from<br>263<br>fewer to<br>252<br>more) | VERY LOW       | CRITICAL       |
| Depres      | sion symptom         | s at endpo                       | oint (follow-up m           | iean 6 weeks; i            | measured witl                | h: CES-D change             | score; Better                     | indicated b                        | y lower va                      | alues)  |                |                |
| 1           | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency    | no serious<br>indirectness | very<br>serious <sup>4</sup> | none                        | 10                                | 12                                 | -                               | SMD<br>0.11<br>higher<br>(0.73<br>lower to                          | VERY LOW       | IMPORTA<br>NT  |

| Quality              | assessment           |                      |                             |                            |                      |                             | No of patien                      | its                                | Effect                      |  |                  |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------------------|------------------------------------|-----------------------------|--|------------------|----------------|
| No of<br>studi<br>es | Design               | Risk of bias         | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Supportiv<br>e<br>counsellin<br>g | No<br>treatme<br>nt or<br>waitlist | Relativ<br>e<br>(95%<br>CI) | Absolut<br>e   | Quality          | Importanc<br>e |
|                      |                      |                      |                             |                            |                      |                             |                                   |                                    |                             | 0.95<br>higher)  |                  |                |
| <b>Depres</b>        | sion symptom         | is at 3-mor          | nth follow-up (fo           | llow-up mean               | 3 months; me         | asured with: CES            | -D/MINI:Depre                     | ession char                        | ige score;                  | <b>Better indi</b>   | cated by lower   | values)        |
| 2                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>5</sup> | reporting bias <sup>3</sup> | 34                                | 40                                 | -                           | SMD 0.7<br>lower<br>(1.17 to<br>0.22<br>lower)               | VERY LOW         | IMPORTA<br>NT  |
| <b>Depres</b>        | sion symptom         | is at 6-mor          | nth follow-up (fo           | llow-up mean               | 6 months; me         | asured with: MIN            | :Depression                       | change sco                         | re; Better                  | indicated b  | y lower values   | )              |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 23                                | 28                                 | -                           | SMD<br>0.47<br>lower<br>(1.03<br>lower to<br>0.09<br>higher) | VERY LOW         | IMPORTA<br>NT  |
| Depres               | sion symptom         | is at 12-mo          | onth follow-up (f           | ollow-up mear              | 12 months; n         | neasured with: M            | INI:Depressio                     | n change s                         | core; Bett                  | er indicated   | d by lower valu  | es)            |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 23                                | 28                                 | -                           | SMD<br>0.34<br>lower<br>(0.9<br>lower to<br>0.21<br>higher)  | VERY LOW         | IMPORTA<br>NT  |
| Function values      |                      | nt at 3-mor          | nth follow-up (fo           | llow-up mean               | 3 months; me         | asured with: CAP            | S: Functional                     | impairmen                          | t; change                   | score; Bett  | ter indicated by | lower          |
| 1                    | randomised<br>trials | no<br>serious        | no serious inconsistency    | no serious indirectness    | serious <sup>5</sup> | reporting bias <sup>3</sup> | 24                                | 28                                 | -                           | SMD<br>0.91<br>lower   | LOW              | IMPORTA<br>NT  |

| Quality              | assessment           |                                  |                             |                            |                      |                             | No of patier                      | its                                | Effect                          |  |                  |               |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|----------------------|-----------------------------|-----------------------------------|------------------------------------|---------------------------------|--|------------------|---------------|
| No of<br>studi<br>es | Design               | Risk of bias                     | Inconsistenc<br>y           | Indirectnes<br>s           | Imprecisio<br>n      | Other consideration s       | Supportiv<br>e<br>counsellin<br>g | No<br>treatme<br>nt or<br>waitlist | Relativ<br>e<br>(95%<br>CI)     | Absolut<br>e   | Quality          | Importance    |
|                      |                      | risk of<br>bias                  |                             |                            |                      |                             |                                   |                                    |                                 | (1.49 to<br>0.34<br>lower)                                   |                  |               |
| Function values      | •                    | nt at 6-mo                       | nth follow-up (fo           | llow-up mean (             | 6 months; me         | asured with: CAP            | S: Functional                     | impairmen                          | it; change                      | score; Bett  | ter indicated by | lower         |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 23                                | 28                                 | -                               | SMD<br>0.44<br>lower (1<br>lower to<br>0.12<br>higher)       | LOW              | IMPORTA<br>NT |
| Function values      | •                    | nt at 12-m                       | onth follow-up (f           | ollow-up mean              | 12 months; n         | neasured with: C            | APS: Function                     | nal impairm                        | ent; chanç                      | ge score; B  | etter indicated  | by lower      |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 23                                | 28                                 | -                               | SMD<br>0.27<br>lower<br>(0.82<br>lower to<br>0.28<br>higher) | LOW              | IMPORTA<br>NT |
| Discon               | tinuation (follo     | ow-up 3-6                        | weeks; assessed             | d with: Number             | of participan        | ts lost to follow-u         | ip for any rea                    | son)                               |                                 |  |                  |               |
| 2                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious inconsistency    | no serious indirectness    | serious <sup>6</sup> | none                        | 6/40<br>(15%)                     | 0/40<br>(0%)                       | RR<br>6.75<br>(0.86 to<br>52.7) | -  | MODERATE         | CRITICAL      |

CAPS= Clinician Administered PTSD Symptom; CES-D= Centre for Epidemiological Studies-Depression; CI=confidence interval; CRIES= Children's Revised Impact of Event Scale; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

### Psychological: Parent training/family intervention

Parent training (CBT with parent-only) versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality              | assessment           |                      |                             |                            |                      |                             | No of patien  | its     | Effect                      |   |             |                |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|----------------------|-----------------------------|---|---------|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias      | Inconsistency               | Indirectness               | Imprecisio<br>n      | Other considerations        | Parent<br>training<br>(CBT with<br>parent-<br>only) | TA<br>U | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| PTSD s               | ymptomatology        | / clinician-         | rated at endpoint           | (follow-up mean            | 12 weeks; me         | easured with: K-SA          | DS-E: PTSD;   | chang   | ge score; l                 | Better indicated                                    | by lower    | values)        |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20  | 14      | -                           | SMD 0.59<br>lower (1.29<br>lower to 0.11<br>higher) | VERY<br>LOW | CRITICAL       |
| PTSD syvalues)       |                      | / clinician-         | rated at 3-month f          | ollow-up (follow           | -up mean 3 m         | onths; measured v           | vith: K-SADS-                                       | E: PT   | SD; chang                   | e score; Better                                     | indicated   | by lower       |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20  | 14      | -                           | SMD 0.63<br>lower (1.33<br>lower to 0.07<br>higher) | VERY<br>LOW | CRITICAL       |
| PTSD syvalues)       |                      | / clinician-         | rated at 6-month f          | ollow-up (follow           | -up mean 6 m         | onths; measured v           | vith: K-SADS-                                       | E: PT   | SD; chang                   | e score; Better                                     | indicated   | by lower       |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious indirectness    | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20  | 14      | -                           | SMD 0.58<br>lower (1.28<br>lower to 0.12<br>higher) | VERY<br>LOW | CRITICAL       |

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>5</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>6</sup> 95% CI crosses both line of no effect and threshold for clinically important harm

| Quality              | assessment           |                      |                             |                            |                              |                             | No of patier  | nts     | Effect                      |   |             |               |
|----------------------|----------------------|----------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|---|---------|-----------------------------|---|-------------|---------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias      | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other considerations        | Parent<br>training<br>(CBT with<br>parent-<br>only) | TA<br>U | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importan      |
| PTSD s<br>values)    | ymptomatology        | y clinician-         | rated at 12-month           | follow-up (follow          | w-up mean 12                 | months; measure             | d with: K-SAD                                       | S-E: F  | PTSD; cha                   | nge score; Bett                                     | er indicat  | ed by lowe    |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 20  | 14      | -                           | SMD 0.42<br>lower (1.11<br>lower to 0.27<br>higher) | VERY<br>LOW | CRITICAL      |
| PTSD s<br>values)    | ymptomatology        | y clinician-         | rated at 2-year foll        | low-up (follow-u           | p mean 2 yeaı                | rs; measured with:          | K-SADS-E: P   | TSD;    | change so                   | ore; Better indi                                    | cated by I  | ower          |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup>         | reporting bias <sup>3</sup> | 20  | 14      | -                           | SMD 0.89<br>lower (1.6 to<br>0.17 lower)            | VERY<br>LOW | CRITICAL      |
| Emotion              |                      | oural probl          | ems-Externalizing           | at endpoint (fol           | low-up mean                  | 12 weeks; measur            | ed with: CBCI                                       | _: Exte | ernalizing;                 | change score;                                       | Better inc  | dicated by    |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 18  | 12      | -                           | SMD 0.63<br>lower (1.38<br>lower to 0.12<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
|                      | nal and behavio      |                      | ems-Externalizing           | at 3-month follo           | ow-up (follow-               | up mean 3 months            | s; measured w                                       | /ith: C | BCL: Exte                   | rnalizing; chan                                     | ge score;   | Better        |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 18  | 12      | -                           | SMD 0.23<br>lower (0.96<br>lower to 0.5<br>higher)  | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment                    |                      |                             |                            |                              |                             | No of patier  | nts     | Effect                      |   |             |               |
|----------------------|-------------------------------|----------------------|-----------------------------|----------------------------|------------------------------|-----------------------------|---|---------|-----------------------------|---|-------------|---------------|
| No of<br>studie<br>s | Design                        | Risk of<br>bias      | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other considerations        | Parent<br>training<br>(CBT with<br>parent-<br>only) | TA<br>U | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importance    |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 18  | 12      | -                           | SMD 0.18<br>lower (0.91<br>lower to 0.55<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
|                      | nal and behavio               |                      | ems-Externalizing           | at 12-month fol            | low-up (follow               | v-up mean 12 mon            | ths; measured                                       | d with: | CBCL: Ex                    | ternalizing; cha                                    | nge scor    | e; Better     |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>5</sup> | reporting bias <sup>3</sup> | 18  | 12      | -                           | SMD 0.07<br>lower (0.8<br>lower to 0.66<br>higher)  | VERY<br>LOW | IMPORTA<br>NT |
|                      | nal and behavior<br>r values) | oural probl          | ems-Externalizing           | at 2-year follow           | -up (follow-up               | mean 2 years; me            | easured with:                                       | CBCL    | : Externali                 | izing; change so                                    | ore; Bett   | er indicated  |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious<br>inconsistency | no serious<br>indirectness | serious <sup>4</sup>         | reporting bias <sup>3</sup> | 18  | 12      | -                           | SMD 0.92<br>lower (1.69 to<br>0.15 lower)           | VERY<br>LOW | IMPORTA<br>NT |
| Depress              | sion symptoms                 | at endpoir           | nt (follow-up mear          | n 12 weeks; mea            | sured with: C                | DI change score; E          | Better indicate                                     | d by I  | ower valu                   | es)   |             |               |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>4</sup>         | reporting bias <sup>3</sup> | 20  | 15      | -                           | SMD 0.86<br>lower (1.56 to<br>0.15 lower)           | VERY<br>LOW | IMPORTA<br>NT |
| Depress              | sion symptoms                 | at 3-mont            | h follow-up (follov         | v-up mean 3 mo             | nths; measure                | ed with: CDI chang          | e score; Bette                                      | er indi | cated by lo                 | ower values)  |             |               |
| 1                    | randomised<br>trials          | serious <sup>1</sup> | no serious inconsistency    | no serious indirectness    | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 20  | 15      | -                           | SMD 0.45<br>lower (1.13<br>lower to 0.23<br>higher) | VERY<br>LOW | IMPORTA<br>NT |

| Quality              | assessment           |                      |                          |                         |                      |                             | No of patien  | ts      | Effect                      |  |             |                |
|----------------------|----------------------|----------------------|--------------------------|-------------------------|----------------------|-----------------------------|---|---------|-----------------------------|--|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of bias         | Inconsistency            | Indirectness            | Imprecisio<br>n      | Other considerations        | Parent<br>training<br>(CBT with<br>parent-<br>only) | TA<br>U | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                    | randomised trials    | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20  | 15      | -                           | SMD 0.32<br>lower (1 lower<br>to 0.35<br>higher)   | VERY<br>LOW | IMPORTA<br>NT  |
| Depress              | ion symptoms         | at 12-mon            | th follow-up (follo      | w-up mean 12 m          | onths; measu         | red with: CDI char          | nge score; Be                                       | tter in | dicated by                  | y lower values)                                    |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | reporting bias <sup>3</sup> | 20  | 15      | -                           | SMD 0.5<br>lower (1.18<br>lower to 0.18<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Depress              | ion symptoms         | at 2-year f          | ollow-up (follow-u       | p mean 2 years;         | measured wi          | th: CDI change sc           | ore; Better inc                                     | licated | d by lower                  | values)  |             |                |
| 1                    | randomised<br>trials | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>4</sup> | reporting bias <sup>3</sup> | 20  | 15      | -                           | SMD 0.86<br>lower (1.56 to<br>0.15 lower)          | VERY<br>LOW | IMPORTA<br>NT  |

CBCL=Children's Behavioural Checklist; CBT=cognitive behavioural therapy; CDI=Children's Depression Inventory; CI=confidence interval; K-SADS=Kiddele Schedulae for Affective Disorder and Schizophrenia; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; TAU=treatment as usual

<sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>2 95%</sup> CI crosses both line of no effect and threshold for clinically important benefit

<sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>4</sup> OIS not met (N<400)

<sup>5 95%</sup> CI crosses line of no effect and thresholds for both clinically important benefit and harm

Parent training + trauma-focused CBT (for child) versus trauma-focused CBT (for child) only for delayed treatment (>3 months) of clinically important sympotms/PTSD

| Quality        | assessment           |                 |                             |                            |                  |                       | No of patie                                       | ents   | Effect                      |   |             |               |
|----------------|----------------------|-----------------|-----------------------------|----------------------------|------------------|-----------------------|---|--|-----------------------------|---|-------------|---------------|
| No of studie s | Design               | Risk of<br>bias | Inconsistency               | Indirectness               | Imprecisio<br>n  | Other consideration s | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importance    |
| PTSD s         | ymptomatolog         | y clinician-    | rated at endpoint           | (follow-up mea             | an 20 weeks; ı   | measured with: A      | DIS-C: PTSD                                       | ; change sc                                  | ore; Better                 | indicated by  | lower val   | ues)          |
| 1              | randomised<br>trials | serious1        | no serious<br>inconsistency | no serious<br>indirectness | serious2         | reporting bias3       | 12  | 12   | -                           | SMD 0.36<br>lower (1.16<br>lower to<br>0.45<br>higher)  | VERY<br>LOW | CRITICAL      |
| PTSD s values) | ymptomatolog         | y clinician-    | rated at 3-month            | follow-up (follo           | w-up mean 3      | months; measure       | d with: ADIS                                      |  | hange sco                   |   | icated by   |               |
| 1              | randomised<br>trials | serious1        | no serious<br>inconsistency | no serious<br>indirectness | serious2         | reporting bias3       | 12  | 12   | -                           | SMD 0.48<br>lower (1.29<br>lower to<br>0.34<br>higher)  | VERY<br>LOW | CRITICAL      |
| <b>Anxiety</b> | symptoms at          | endpoint (f     | ollow-up mean 20            | ) weeks; measu             | red with: RCI    | MAS; change scor      | e; Better ind                                     | licated by lo                                | wer values                  | 5)  |             |               |
| 1              | randomised<br>trials | serious1        | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3       | 12  | 12   | -                           | SMD 0.14<br>higher<br>(0.66 lower<br>to 0.94<br>higher) | VERY<br>LOW | IMPORTA<br>NT |
| <b>Anxiety</b> | symptoms at          | 3-month fo      | llow-up (follow-u           | p mean 3 montl             | ns; measured     | with: RCMAS; ch       | ange score;                                       | Better indica                                | ated by low                 | ver values)   |             |               |
| 1              | randomised<br>trials | serious1        | no serious<br>inconsistency | no serious indirectness    | very<br>serious4 | reporting bias3       | 12  | 12   | -                           | SMD 0.03<br>higher<br>(0.77 lower<br>to 0.83<br>higher) | VERY<br>LOW | IMPORTA<br>NT |

| Quality             | assessment                              |                 |                             |                            |                  |                                      | No of patie                                       | ents   | Effect                      |  |             |                |
|---------------------|---|-----------------|-----------------------------|----------------------------|------------------|--------------------------------------|---|--|-----------------------------|--|-------------|----------------|
| No of studie s      | Design                                  | Risk of<br>bias | Inconsistency               | Indirectness               | Imprecisio<br>n  | Other consideration s                | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute   | Qualit<br>y | Importanc<br>e |
| 1                   | randomised<br>trials                    | serious1        | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3                      | 12  | 12   | -                           | SMD 0.29<br>lower (1.09<br>lower to<br>0.52<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| 1                   | randomised<br>trials                    | serious1        | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | red with: CDI; chare reporting bias3 | 12  | 12   |                             | SMD 0.07<br>lower (0.87<br>lower to<br>0.73<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Emotion lower value |   | oural probl     | ems-Internalizing           | g at endpoint (fo          | ollow-up mear    | n 20 weeks; meas                     | ured with: Cl                                     | BCL: Interna                                 | llizing; cha                | inge score; Be   | etter indi  | cated by       |
| 1                   | randomised<br>trials                    | serious1        | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3                      | 12  | 12   | -                           | SMD 0.29<br>lower (1.1<br>lower to<br>0.51<br>higher)  | VERY<br>LOW | IMPORTA<br>NT  |
|                     |   |                 | ems-Internalizing           | at 3-month fol             | low-up (follow   | /-up mean 3 mont                     | hs; measure                                       | d with: CBC                                  | L: Internal                 | izing; change  | score; B    | etter          |
| 1                   | ed by lower val<br>randomised<br>trials | serious1        | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3                      | 12  | 12   | -                           | SMD 0.15<br>lower (0.95<br>lower to<br>0.66<br>higher) | VERY<br>LOW | IMPORTA<br>NT  |
| Emotion lower value |   | oural probl     | ems-Externalizin            | g at endpoint (f           | ollow-up mea     | n 20 weeks; meas                     | ured with: C                                      | BCL: Extern                                  | alizing; ch                 | nange score; I   | Better inc  | licated by     |
| 1                   | randomised<br>trials                    | serious1        | no serious<br>inconsistency | no serious indirectness    | serious2         | reporting bias3                      | 12  | 12   | -                           | SMD 0.79<br>lower (1.63<br>lower to                    | VERY<br>LOW | IMPORTA<br>NT  |

| Quality              | assessment           |                                  |                             |                            |                  |                       | No of patie                                       | ents   | Effect                      |   |             |                |
|----------------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------|-----------------------|---|--|-----------------------------|---|-------------|----------------|
| No of<br>studie<br>s | Design               | Risk of<br>bias                  | Inconsistency               | Indirectness               | Imprecisio<br>n  | Other consideration s | Trauma-<br>focused<br>CBT +<br>parent<br>training | Trauma-<br>focused<br>CBT<br>(child<br>only) | Relativ<br>e<br>(95%<br>CI) | Absolute  | Qualit<br>y | Importanc<br>e |
| Emotio               | nal and hohavi       | oural probl                      | ome-Eytornalizin            | g at 3-month fo            | llow-up (follo   | w-up mean 3 mon       | the measur  | ed with: CR                                  | CI · Evtern                 | 0.04<br>higher)   | e score.    | Retter         |
|                      | d by lower val       |                                  | ems-Externanzm              | g at 3-month to            | now-up (rono     |                       | uis, illeasui                                     | eu with. Obt                                 | JE. EXIGIII                 | anzing, chang   | e score,    | Detter         |
| 1                    | randomised<br>trials | serious1                         | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3       | 12  | 12   | -                           | SMD 0.14<br>lower (0.94<br>lower to<br>0.67<br>higher)    | VERY<br>LOW | IMPORTA<br>NT  |
| Global f             |                      |                                  | ollow-up mean 20            | ) weeks; measu             | red with: GAF    | ; change score;       |   |  | er values)                  |   |             |                |
| 1                    | randomised<br>trials | serious1                         | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | reporting bias3       | 12  | 12   | -                           | SMD 0.3<br>higher (0.5<br>lower to<br>1.11<br>higher)     | VERY<br>LOW | IMPORTA<br>NT  |
| Global f             | unctioning at        | 3-month fo                       | llow-up (follow-u           | p mean 3 month             | ns; measured     | with: GAF; chang      |   |  | d by higher                 |   |             |                |
| 1                    | randomised<br>trials | serious1                         | no serious<br>inconsistency | no serious<br>indirectness | serious2         | reporting bias3       | 12  | 12   | -                           | SMD 0.66<br>higher<br>(0.16 lower<br>to 1.49<br>higher)   | VERY<br>LOW | IMPORTA<br>NT  |
|                      |                      |                                  |                             |                            |                  | ants lost to follow   |   |  |                             |   |             |                |
| 1                    | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious4 | none                  | 3/12<br>(25%)                                     | 3/12<br>(25%)                                | RR 1<br>(0.25 to<br>4)      | 0 fewer per<br>1000 (from<br>188 fewer<br>to 750<br>more) | LOW         | CRITICAL       |

ADIS-C= Anxiety Disorder Interview Schedule-Child version; CBCL= Child Behavioural Checklist; CBT=cognitive behavioural therapy; CDI= Children's Depression Inventory; CI=confidence interval; GAF= Global Assessment of Functioning; PTSD=post-traumatic stress disorder; RCMAS=; RR=risk ratio; SMD=standardised mean difference

Family therapy versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

|                | assessment           |                                  | 1                        |                            |                      |                       | No of pa              |                    | Effect                       |  |          |                |
|----------------|----------------------|----------------------------------|--------------------------|----------------------------|----------------------|-----------------------|-----------------------|--------------------|------------------------------|--|----------|----------------|
| No of studies  | Design               | Risk of bias                     | Inconsistency            | Indirectness               | Imprecisio<br>n      | Other consideration s | Family<br>therap<br>y | Waitli<br>st       | Relative<br>(95%<br>CI)      | Absolute   | Quality  | Importanc<br>e |
| PTSD s         | ymptomatolog         | y (follow-u                      | p mean 0.1 weeks         | s; measured wi             | th: UCLA PTS         | D-RI; change sco      | re; Better            | indicated          | l by lower v                 | /alues)  |          |                |
| 1              | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency | no serious<br>indirectness | serious <sup>2</sup> | none                  | 75                    | 74                 | -                            | SMD 0.37<br>lower (0.7<br>to 0.05<br>lower)              | LOW      | CRITICAL       |
| <b>Anxiety</b> | symptoms (fo         | llow-up me                       | ean 0.1 weeks; m         | easured with: R            | CMAS; T-sco          | res change score      | ; Better in           | dicated b          | y lower val                  | ues)   |          |                |
| 1              | randomised<br>trials | serious <sup>1</sup>             | no serious inconsistency | no serious indirectness    | serious <sup>2</sup> | none                  | 75                    | 74                 | -                            | SMD 0.09<br>higher<br>(0.24 lower<br>to 0.41<br>higher)  | LOW      | IMPORTA<br>NT  |
| Discon         | tinuation (follo     | w-up mean                        | 0.1 weeks; asses         | ssed with: Num             | ber of particip      | ants lost to follow   | w-up for a            | ny reasoi          | n)                           |  |          |                |
| 1              | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious inconsistency | no serious indirectness    | serious <sup>3</sup> | none                  | 29/76<br>(38.2%<br>)  | 5/74<br>(6.8%<br>) | RR 5.65<br>(2.31 to<br>13.8) | 314 more<br>per 1000<br>(from 89<br>more to<br>865 more) | MODERATE | CRITICAL       |

CI=confidence interval; PTSD=post-traumatic stress disorder; RCMAS=Revised Children Manifest Anxiety Scale; RR=risk ratio; SMD=standardised mean difference; UCLA PTSD-RI=UCLA PTSD-Reaction Index;

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> 95% CI crosses both line of no effect and threshold for clinically important benefit

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> OIS not met (events<300)

### **Psychological: Play therapy**

Play therapy versus TAU for the delayed treatment (>3 months) of clinically important symptoms/PTSD Non-directive counselling

| Quality        | assessment           |                              |                          |                         |                              |                             | No of pa            | tients             | Effect                       |   |             |                |
|----------------|----------------------|------------------------------|--------------------------|-------------------------|------------------------------|-----------------------------|---------------------|--------------------|------------------------------|---|-------------|----------------|
| No of studie s | Design               | Risk of bias                 | Inconsistency            | Indirectness            | Imprecisio<br>n              | Other considerations        | Play<br>therap<br>y | TAU                | Relative<br>(95% CI)         | Absolute  | Qualit<br>y | Importanc<br>e |
| PTSD s         | ymptomatology        | self-rated                   | (follow-up mean          | 3 weeks; measu          | red with: CRIE               | S change score; I           | Better indi         | cated by           | / lower valu                 | es)   |             |                |
| 1              | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 69                  | 60                 | -                            | SMD 1.07<br>lower (1.44 to<br>0.7 lower)              | VERY<br>LOW | CRITICAL       |
| <b>Anxiety</b> | symptoms (fol        | low-up mea                   | an 3 weeks; meas         | ured with: SCAS         | S; change sco                | re; Better indicate         | d by lowe           | r values)          |                              |   |             |                |
| 1              | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 69                  | 60                 | -                            | SMD 1.87<br>lower (2.29 to<br>1.45 lower)             | VERY<br>LOW | IMPORTA<br>NT  |
| Depress        | sion symptoms        | (follow-up                   | mean 3 weeks; m          | easured with: S         | MFQ; change                  | score; Better indi          | cated by l          | ower val           | ues)                         |   |             |                |
| 1              | randomised<br>trials | very<br>serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup>         | reporting bias <sup>3</sup> | 69                  | 60                 | -                            | SMD 1.34<br>lower (1.73 to<br>0.96 lower)             | VERY<br>LOW | IMPORTA<br>NT  |
| Discont        | inuation (follow     | v-up mean                    | 3 weeks; assesse         | d with: Number          | of participants              | s lost to follow-up         | for any re          | ason)              |                              |   |             |                |
| 1              | randomised<br>trials | serious <sup>1</sup>         | no serious inconsistency | no serious indirectness | very<br>serious <sup>4</sup> | none                        | 0/69<br>(0%)        | 2/62<br>(3.2<br>%) | RR 0.18<br>(0.01 to<br>3.68) | 26 fewer per<br>1000 (from 32<br>fewer to 86<br>more) | VERY<br>LOW | CRITICAL       |

Cl=confidence interval; CRIES; PTSD=post-traumatic stress disorder; RR=risk ratio; SCAS= Spence Children's Anxiety Scale; SMD=standardised mean difference; SMFQ=Short Mood and Feeling Questionnaires; TAU=treatment as usual

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains

<sup>&</sup>lt;sup>2</sup> OIS not met (N<400)

<sup>&</sup>lt;sup>3</sup> Data is not reported/cannot be extracted for all outcomes

<sup>&</sup>lt;sup>4</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

### Play therapy versus trauma-focused CBT for the delayed treatment (>3 months) of clinically important symptoms/PTSD

|                | assessment           |                                  |                          |                            |                              | ·                    | No of pa            |                           | Effect                       |   |             |                |
|----------------|----------------------|----------------------------------|--------------------------|----------------------------|------------------------------|----------------------|---------------------|---------------------------|------------------------------|---|-------------|----------------|
| No of studie s | Design               | Risk of bias                     | Inconsistency            | Indirectness               | Imprecisio<br>n              | Other considerations | Play<br>therap<br>y | Trauma-<br>focused<br>CBT | Relative<br>(95% CI)         | Absolute  | Qualit<br>y | Importanc<br>e |
| PTSD s         | ymptomatolog         | y self-rated                     | (follow-up mean          | 12 weeks; meas             | sured with: UC               | LA PTSD-RI; char     | nge score           | ; Better indi             | icated by lo                 | wer values)   |             |                |
| 1              | randomised trials    | serious <sup>1</sup>             | no serious inconsistency | no serious<br>indirectness | very<br>serious <sup>2</sup> | none                 | 14                  | 12                        | -                            | SMD 0.11<br>lower (0.88<br>lower to<br>0.66 higher)         | VERY<br>LOW | CRITICAL       |
| Discont        | inuation (follo      | w-up mean                        | 12 weeks; assess         | ed with: Number            | er of participa              | nts lost to follow-u | up for any          | reason)                   |                              |   |             |                |
| 1              | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious inconsistency | no serious indirectness    | very<br>serious <sup>2</sup> | none                 | 0/14 (0%)           | 5/17<br>(29.4%)           | RR 0.11<br>(0.01 to<br>1.82) | 262 fewer<br>per 1000<br>(from 291<br>fewer to 241<br>more) | LOW         | CRITICAL       |

CI=confidence interval; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference; TAU=treatment as usual; UCLA PTSD-RI=UCLA PTSD-Reaction Index

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains <sup>2</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

### **Psychosocial: Art therapy**

Art therapy (+ TAU) versus attention-placebo (+ TAU) for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality        | Quality assessment   |                                  |                          |                         |                      |                             | No of pat                 | ients                            | Effect                      |  |             |                |
|----------------|--|----------------------------------|--------------------------|-------------------------|----------------------|-----------------------------|---------------------------|----------------------------------|-----------------------------|--|-------------|----------------|
| No of studie s | Design   | Risk of bias                     | Inconsistency            | Indirectness            | Imprecisio<br>n      | Other considerations        | Art<br>therapy<br>(+ TAU) | Attention-<br>placebo (+<br>TAU) | Relativ<br>e<br>(95%<br>CI) | Absolute                                     | Quali<br>ty | Importanc<br>e |
|                | PTSD symptomatology clinician-rated (follow-up mean 16 weeks; measured with: UCLA PTSD-RI administered via structured intervindicated by lower values) |                                  |                          |                         |                      |                             |                           |                                  |                             |  | hange s     | core; Better   |
| 1              | randomised<br>trials   | no<br>serious<br>risk of<br>bias | no serious inconsistency | no serious indirectness | serious <sup>1</sup> | reporting bias <sup>2</sup> | 14                        | 15                               | -                           | SMD 1.79<br>lower (2.67<br>to 0.91<br>lower) | LOW         | CRITICAL       |

Cl=confidence interval; PTSD=post-traumatic stress disorder; SMD=standardised mean difference; TAU=treatment as usual; UCLA PTSD-RI= UCLA PTSD-Reaction Index

OIS not met (N<400)

### **Psychosocial: Meditation**

Meditation versus waitlist for the delayed treatment (>3 months) of clinically important symptoms/PTSD

| Quality        | quality assessment |                      |                          |                         |                      | No of patients       |                | Effect       |                      |                         |             |                |
|----------------|--------------------|----------------------|--------------------------|-------------------------|----------------------|----------------------|----------------|--------------|----------------------|-------------------------|-------------|----------------|
| No of studie s | Design             | Risk of bias         | Inconsistency            | Indirectness            | Imprecisio<br>n      | Other considerations | Meditatio<br>n | Waitli<br>st | Relative<br>(95% CI) | Absolute                | Quali<br>ty | Importanc<br>e |
| PTSD sy        | ymptomatolog       | y self-rated         | (follow-up mean          | 6 weeks; measu          | red with: HTC        | change score; B      | etter indicate | ed by low    | er values)           |                         |             |                |
| 1              | randomised trials  | serious <sup>1</sup> | no serious inconsistency | no serious indirectness | serious <sup>2</sup> | none                 | 38             | 39           | -                    | SMD 1.65<br>lower (2.17 | LOW         | CRITICAL       |

<sup>&</sup>lt;sup>2</sup> Data is not reported/cannot be extracted for all outcomes and this is interim report but unable to locate full report

| Quality        |                      |                                  |                             |                            |                              |                      | Effect         |                    |                             |   |             |                |
|----------------|----------------------|----------------------------------|-----------------------------|----------------------------|------------------------------|----------------------|----------------|--------------------|-----------------------------|---|-------------|----------------|
| No of studie s | Design               | Risk of bias                     | Inconsistency               | Indirectness               | Imprecisio<br>n              | Other considerations | Meditatio<br>n | Waitli<br>st       | Relative<br>(95% CI)        | Absolute  | Quali<br>ty | Importanc<br>e |
| Discont        | inuation (follo      | w-up mean                        | 6 weeks: assesse            | d with: Number             | of participant               | s lost to follow-up  | o for any reas | son)               |                             | to 1.13<br>lower)                                     |             |                |
| 1              | randomised<br>trials | no<br>serious<br>risk of<br>bias | no serious<br>inconsistency | no serious<br>indirectness | very<br>serious <sup>3</sup> | none                 | 3/41<br>(7.3%) | 2/41<br>(4.9%<br>) | RR 1.5<br>(0.26 to<br>8.51) | 24 more per<br>1000 (from<br>36 fewer to<br>366 more) | LOW         | CRITICAL       |

CI=confidence interval; HTQ= Harvard Trauma Questionnaire; PTSD=post-traumatic stress disorder; RR=risk ratio; SMD=standardised mean difference

<sup>&</sup>lt;sup>1</sup> Risk of bias is high or unclear across multiple domains
<sup>2</sup> OIS not met (N<400)
<sup>3</sup> 95% CI crosses line of no effect and thresholds for both clinically important benefit and harm

# Appendix G- Health economic evidence study selection

Health economic evidence study selection for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

A global health economics search was undertaken for all areas covered in the guideline. The flow diagram of economic article selection across all reviews is provided in Appendix A of Supplement 1 – Methods Chapter'.

## **Appendix H- Economic evidence tables**

Health economic evidence tables for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

### Psychological interventions - references to included studies

Gospodarevskaya E and Segal L (2012) Cost-utility analysis of different treatments for post-traumatic stress disorder in sexually abused children. Child and Adolescent Psychiatry and Mental Health 2012; 6(15)

McCrone P, Weeramanthri T, Knapp MG et al. (2005) Cost-Effectiveness of Individual versus Group Psychotherapy for Sexually Abused Girls. Child and Adolescent Mental Health 10(1), 26-31

Mihalopoulos C, Magnus A, Lal A et al. (2015) Is implementation of the 2013 Australian treatment guidelines for posttraumatic stress disorder cost-effective compared to current practice? A cost-utility analysis using QALYs and DALYs. Australian and New Zealand Journal of Psychiatry 49(4), 360-376

Shearer J, Papanikolaou N, Meiser-Stedman R et al. (2018). Cost-effectiveness of cognitive therapy as an early intervention for post-traumatic stress disorder in children and adolescents: a trial based evaluation and model. Journal of Child Psychology and Psychiatry, doi: 10.1111/jcpp.12851

| Study<br>Country<br>Study type  | Intervention details  | Study population<br>Study design<br>Data sources   | Costs and outcomes: description and values   | Results: Cost-<br>effectiveness   | Comments  |
|---|---|--|--|---|---|
| Gospodarev<br>skaya &<br>Segal, 2012<br>Australia<br>Cost-utility<br>analysis | Interventions: Trauma-focused cognitive behavioural therapy, comprising 12 weekly sessions of 45 min each (TF-CBT)  TF-CBT combined with selective serotonin reuptake inhibitors (SSRIs)  Non-directive supportive counselling comprising 12 weekly sessions of 45 min each  No treatment | 10-year-old children who met all or most of the PTSD diagnostic criteria, including at least one symptom of avoidance or re-experiencing, who were eligible for TF-CBT; some of the children had comorbid depression  Decision-analytic economic modelling  Source of efficacy data: meta-analyses of trials and indirect comparisons  Source of resource use data: published trial data  Source of unit costs: national sources | Costs: therapists' time [psychologist, psychiatrist, GP, social worker], medication, parental group or psychoeducational sessions; beyond 12 months: medication + GP costs  Total cost per child (12 months): TF-CBT \$2051 TF-CBT + SSRI £2226 Counselling \$2074 No treatment 0  Total cost per child (31 years): TF-CBT \$2096 TF-CBT + SSRI \$2270 Counselling \$2123 No treatment 0  Outcome measure: QALY based on the Assessment of Quality of Life measure | At 12 months: Counselling dominated by TF-CBT  ICERS TF-CBT + SSRI vs TF-CBT \$17,520/QALY  TF-CBT vs no treatment \$22,790/QALY  At 31 years: Counselling dominated by TF-CBT  ICERS TF-CBT + SSRI vs TF-CBT \$2,901/QALY  TF-CBT vs no treatment \$1,650/QALY  Results sensitive to variation in clinical effectiveness | Perspective: mental health system Currency: Aus\$ Cost year: 2011 Time horizon: 31 years Discounting: 5% Applicability: partially applicable Quality: potentially serious limitations |

| Study<br>Country<br>Study type | Intervention details | Study population Study design Data sources | Costs and outcomes: description and values   | Results: Cost-<br>effectiveness | Comments |
|--------------------------------|----------------------|--|--|---------------------------------|----------|
|                                |                      |  | (AQoL-4D), Australian values used  Total QALYs per child (12 months): TF-CBT 0.96 TF-CBT + SSRI 0.97 Counselling 0.93 No treatment 0.87  Total QALYs per child (31 years): TF-CBT 12.86 TF-CBT + SSRI 12.92 Counselling 12.61 No treatment 11.59 |                                 |          |

| Study<br>Country<br>Study type                       | Intervention details   | Study population Study design Data sources  | Costs and outcomes: description and values   | Results: Cost-<br>effectiveness  | Comments   |
|--|--|---|--|--|--|
| McCrone et al., 2005  UK  Cost-consequenc e analysis | Interventions: Individual psychotherapy comprising up to 30 sessions of focused psychoanalytical psychotherapy  Group psychoeducational therapy comprising up to 18 sessions with psychotherapeutic and psychoeducational components delivered to groups of 5 girls  both interventions included carers' support | Sexually abused girls (6-14 years old) with symptoms of emotional or behavioural disturbance, 73% of whom had PTSD  RCT (Trowel 2002)  Source of efficacy and resource use data: RCT (N=75; at 1-year follow up: n=58; at 2-year follow up: n=54)  Source of unit costs: national sources | Costs: intervention: therapists' time, including introductory meeting, initial assessment, therapy, carers' support, supervision of therapists and carers' workers, follow up  Mean cost per child: Individual therapy £3195 Group therapy £1949; p <0.001  Outcome measures: global impairment of functioning measured using the K-GAS; Orvaschel's PTSD scale  Outcomes: difference in improvements in global impairment of functioning between interventions not statistically significant; individual therapy showed greater improvements in manifestations of PTSD compared with group therapy; effect size ranging from 0.60 to 0.79 | Individual therapy more costly and more effective in terms of reducing PTSD symptoms | Perspective: providers of mental health services to children and support to parents (intervention costs only) Currency: UK£ Cost year: 1999 Time horizon: 2 years Discounting: NA Applicability: partially applicable Quality: potentially serious limitations |

| Study<br>Country<br>Study type  | Intervention details   | Study population Study design Data sources  | Costs and outcomes: description and values   | Results: Cost-<br>effectiveness  | Comments   |
|---|--|---|--|--|--|
| Mihalopoulo<br>s et al.,<br>2015<br>Australia<br>Cost-utility<br>analysis | Interventions: Trauma-focused cognitive behavioural therapy (TF-CBT) (8-10 individual sessions) delivered by a psychologist  Treatment as usual (TAU): non-evidence-based care comprising consultation with healthcare professionals | Prevalent cases of children and adolescents with PTSD in Australia in 2012, in receipt of non-evidence-based care  Decision-analytic economic modelling  Source of efficacy data: meta-analyses of TF-CBT trials  Source of resource use data: published trial and epidemiological data; expert opinion  Source of unit costs: national sources | Costs: intervention (psychologist, GP)  Mean incremental cost (million) per eligible population (95% CI): TF-CBT vs TAU \$0.87 (\$0.28 to \$2.2)  Primary outcome measure: QALY based on the Assessment of Quality of Life measure (AQoL-4D), Australian values used [DALY also considered]  Mean incremental number of QALYs per eligible population (x1,000) (95% CI): TF-CBT vs TAU 0.09 (0.00 to 0.25) | ICER of TF-CBT vs TAU: \$8900/QALY  Probability of TF-CBT being cost-effective 1.0 at a willingness to pay of \$50,000/QALY  Results most sensitive to PTSD prevalence, effectiveness, adherence and eligibility for CBT | Perspective: health sector (government & service user (intervention costs only) Currency: Aus\$ Cost year: 2012 Time horizon: 5 years Discounting: NA Applicability: partially applicable Quality: potentially serious limitations |

| Study<br>Country<br>Study type                  | Intervention details   | Study population Study design Data sources  | Costs and outcomes: description and values   | Results: Cost-<br>effectiveness  | Comments   |
|---|--|---|--|--|--|
| Shearer et al., 2018  UK  Cost-utility analysis | Interventions: Trauma-focused cognitive therapy (TF-CT) (10 weekly individual sessions) delivered by a trained clinical psychologist  Waitlist | Children and adolescents aged 8-17 years, who had experienced a single traumatic event in the previous 2 to 6 months and met age-appropriate diagnosis of PTSD  RCT (Meiser-Stedman 2010/2017) and decision-analytic economic modelling  Source of efficacy data: RCT (n=29)  Source of resource use data: RCT (n=29)  Source of unit costs: national sources | Costs: intervention (psychologist), inpatient and outpatient care, emergency department, ambulance, community staff (GP, GP nurse, district nurse, paediatrician, clinical psychologist, CAMHS worker, counsellor, educational psychologist), advice service, social services, other services, medication  Mean cost per child: TF-CT: £4,865; WL: £4,768 Difference: £97  Primary outcome measure: QALY based on Strengths and Difficulties Questionnaire (SDQ) ratings, mapped onto the Child Health Utility index 9D (CHU-9D), Australian values used  Mean QALYs per child: TF-CT: 2.370; WL: 2.324 Difference: 0.0577 | ICER of TF-CT vs WL: £2,205/QALY  Probability of TF-CT being cost-effective 0.60-0.69 at a willingness to pay of £20,000-£30,000/QALY, respectively  Completer case analysis: ICER £2,806/QALY; probability of TF-CT being cost-effective: 0.69-0.75 at a willingness to pay of £20,000-£30,000/QALY, respectively  Including psychologist training costs: ICER £16,187/QALY; probability of TF-CT being cost-effective: 0.51-0.62 at a willingness to pay of £20,000-£30,000/QALY, respectively | Perspective: NHS/PSS Currency: UK£ Cost year: 2014 prices Time horizon: 3 years Discounting: 3.5% Applicability: partially applicable Quality: potentially serious limitations |

# **Appendix I – Health economic evidence profiles**

Health economic evidence profiles for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

### **Psychological interventions**

| Economic evidence profile: trauma-focused cognitive behavioural therapy (TF-CBT) versus treatment as usual (TAU) for the treatment of children and young people with PTSD |  |                                      |   |                                       |                    |                                 |  |  |  |
|---|--|--------------------------------------|---|---------------------------------------|--------------------|---------------------------------|--|--|--|
| Study and country   | Limitation s                                       | Applicability                        | Other comments  | Increment<br>al cost (£) <sup>1</sup> | Incremental effect | ICER<br>(£/effect) <sup>1</sup> | Uncertainty <sup>1</sup>   |  |  |
| Mihalopoulo<br>s<br>Australia   | Potentially<br>serious<br>limitations <sup>2</sup> | Partially applicable <sup>3</sup>    | Population: prevalent cases of children and adolescents with PTSD in Australia in 2012, in receipt of non-evidence-based care Outcome: QALY [and DALY]                                  | £0.39<br>million                      | 90                 | £3954                           | Probability of TF-CBT being cost-effective 1.0 at a willingness to pay of £22,214/QALY  Results most sensitive to PTSD prevalence, effectiveness, adherence and eligibility for CBT  |  |  |
| Shearer   | Potentially<br>serious<br>limitations <sup>4</sup> | Partially<br>applicable <sup>5</sup> | Population: Children and adolescents aged 8-17 years, who had experienced a single traumatic event in the previous 2 to 6 months and met ageappropriate diagnosis of PTSD Outcome: QALY | £99                                   | 0.0577             | £2254                           | Probability of TF-CT being cost-effective 0.60-0.69 at a WTP of £20,000-£30,000/QALY, respectively Completer case analysis: ICER £2,869/QALY; probability of TF-CT being cost-effective: 0.69-0.75 at a WTP of £20,000-£30,000/QALY, respectively Including psychologist training costs: |  |  |

Economic evidence profile: trauma-focused cognitive behavioural therapy (TF-CBT) versus treatment as usual (TAU) for the treatment of children and young people with PTSD

ICER £16,549/QALY; probability of TF-CT being cost-effective: 0.51-0.62 at a WTP of £20,000-£30,000/QALY, respectively

- 1. Costs converted and uplifted to 2016 UK pounds using purchasing power parity (PPP) exchange rates and the UK HCHS index (Curtis & Burns, 2016).
- 2. Time horizon 5 years (for benefits, as costs were measured over 10 weeks); analysis based on economic modelling; effectiveness based on meta-analyses of TF-CBT trials; resource use based on trial and epidemiological data and expert opinion; national unit costs used; PSA conducted; consideration of intervention costs only
- 3. Australian study; health sector perspective; QALY estimates based on the Assessment of Quality of Life measure (AQoL-4D, Australian values used)
- 4. Time horizon 3 years; analysis based on a 11-week RCT and extrapolation of findings using economic modelling; effectiveness and resource use based on RCT (n=29); relapse over 3 years assumed to be zero; national unit costs used; PSA conducted
- 5. UK study, NHS/PSS perspective; QALY estimates based on Strengths and Difficulties Questionnaire (SDQ) ratings, which were then mapped onto the Child Health Utility index 9D (CHU-9D, Australian values used)

Economic evidence profile: trauma-focused cognitive behavioural therapy (TF-CBT) alone or in combination with SSRIs versus non-direct counselling versus no treatment for the treatment of children and young people with PTSD

| Study and country                                 | Limitation<br>s                                    | Applicability                        | Other comments   | Increment<br>al cost (£)<br>vs no<br>treatment <sup>1</sup> | Increment<br>al effect vs<br>no<br>treatment           | ICER<br>(£/effect) <sup>1</sup>   | Uncertainty <sup>1</sup>                                 |
|---|--|--------------------------------------|--|---|--|---|--|
| Gospodarev<br>skaya &<br>Segal, 2012<br>Australia | Potentially<br>serious<br>limitations <sup>2</sup> | Partially<br>applicable <sup>3</sup> | Population: 10-year-<br>old children who met<br>all or most of PTSD<br>diagnostic criteria,<br>including at least one<br>symptom of avoidance<br>or re-experiencing,<br>who were eligible for<br>TF-CBT; some of the<br>children had comorbid<br>depression<br>Outcome: QALY | TF-CBT<br>£1042<br>Combo<br>£1128<br>Counselling<br>£1055   | TF-CBT<br>1.27<br>Combo<br>1.33<br>Counselling<br>1.02 | Counselling<br>dominated by<br>TF-CBT<br>Combo vs<br>TF-CBT<br>£1442<br>TF-CBT vs<br>no treatment<br>£820 | Results sensitive to variation in clinical effectiveness |

# Economic evidence profile: trauma-focused cognitive behavioural therapy (TF-CBT) versus treatment as usual (TAU) for the treatment of children and young people with PTSD

- 1. Costs converted and uplifted to 2016 UK pounds using purchasing power parity (PPP) exchange rates and the UK HCHS index (Curtis & Burns, 2016).
- 2. Time horizon 31 years (discounting 5% annually); analysis based on economic modelling; effectiveness based on meta-analyses of trials and indirect comparisons; resource use based on trial data; national unit costs used; PSA conducted; consideration of a narrow range of costs; relapses of PTSD after successful treatment due to the original traumatic event not considered
- 3. Australian study; mental health system perspective; QALY estimates based on the Assessment of Quality of Life measure (AQoL-4D, Australian values used)

Economic evidence profile: Individual short-term psychodynamic psychotherapy versus group psychoeducational therapy for the treatment of children and young people with PTSD

| Study and country            | Limitation<br>s                              | Applicability                     | Other comments  | Increme<br>ntal cost<br>(£) <sup>1</sup> | Increment al effect   | ICER<br>(£/effect) <sup>1</sup> | Uncertainty <sup>1</sup>  |
|------------------------------|--|-----------------------------------|---|--|---|---------------------------------|---|
| McCrone et<br>al, 2005<br>UK | Potentially serious limitations <sup>2</sup> | Partially applicable <sup>3</sup> | Population: Sexually abused girls (6-14 years old) with symptoms of emotional or behavioural disturbance, 73% of whom had PTSD Outcomes: global impairment of functioning measured using the K-GAS; Orvaschel's PTSD scale Costs and outcomes not synthesised | £2051                                    | effect size<br>on PTSD<br>symptoms<br>ranging<br>from 0.60<br>to 0.79 | NA                              | Individual therapy significantly costlier than group therapy.  Difference in improvements in global impairment of functioning between interventions not statistically significant; individual therapy showed greater improvements in manifestations of PTSD compared with group therapy |

- 1. Costs uplifted to 2016 UK pounds using the UK HCHS index (Curtis & Burns, 2016).
- 2. Time horizon 2 years (up to end of intervention); analysis based on RCT (N=75; at 1-year follow up: n=58; at 2-year follow up: n=54); national unit costs used; consideration of intervention costs only; no synthesis of costs and outcomes
- 3. UK study; providers of mental health services perspective; no QALYs estimated

| Economic evidence profile: various psychological interventions for the treatment of children and young people with PTSD |                                   |                                  |                  |  |  |  |   |  |  |
|---|-----------------------------------|----------------------------------|------------------|--|--|--|---|--|--|
| Study and country   | Limitation<br>s                   | Applicability                    | Other comments   | Incremental cost<br>vs no treatment<br>(£) <sup>1</sup>  | Incremental QALY vs no treatment   | ICER (£/effect) <sup>1</sup>   | Uncertainty <sup>1</sup>  |  |  |
| Guideline<br>economic<br>analysis<br>UK   | Minor<br>limitations <sup>2</sup> | Directly applicable <sup>3</sup> | Outcome:<br>QALY | Support counsel 790 Group CBT -315 Cogn process therapy 75 CT -766 Narrative exposure -629 Prolonged exposure 87 EMDR -216 Family therapy 20 Play therapy -286 Parent training -14 | Support counsel 0.06 Group CBT 0.10 Cogn process therapy 0.15 CT 0.35 Narrative exposure 0.20 Prolonged exposure 0.18 EMDR 0.12 Family therapy 0.05 Play therapy 0.18 Parent training 0.12 | CT 44,993 Narrative exposure 41,966 Prolonged exposure 40,742 Play therapy 41,109 Cogn process therapy 40,178 Parent training 39,788 EMDR 39,920 Group CBT 39,687 Support counsel 37,753 Family therapy 38,222 No treatment 37,304 | Prob of cost effectiveness at WTP £20,000/QALY: cogn therapy 0.78; narrative exposure 0.08; play therapy 0.05; prolonged exposure 0.01; cogn process therapy 0.00; EMDR 0.01; parent train 0.03; group CBT 0.01; family therapy 0.01; support counsel; 0.00; no treat 0.01 Results robust to changes in risk of relapse |  |  |

<sup>1.</sup> Costs uplifted to 2017 UK pounds using the UK hospital & community health services (HCHS) index (Curtis & Burns, 2017).

<sup>2.</sup> Decision-analytic hybrid model (decision-tree + Markov); time horizon 3 years; relative effects based on guideline systematic review and NMA; baseline effects & other clinical input parameters derived from published literature and the committee's expert advice; resource use based on RCT data; national unit costs used; PSA conducted; CEACs & CEAF presented

<sup>3.</sup> UK study; NHS & PSS perspective; QALY estimates based on the Assessment of Quality of Life measure (AQoL-4D, Australian values used)

# Appendix J – Health economic analysis: cost effectiveness of interventions for the delayed (>3 months) treatment of PTSD in children and young people

Health economic analysis for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

### Introduction - objective of economic modelling

The choice of treatment for children and young people with PTSD was identified by the committee and the guideline health economist as an area with potentially major resource implications. Existing economic evidence in this area is very limited and does not cover the full range of available interventions for children and young people with PTSD in the UK. On the other hand, clinical evidence was judged to be sufficient and of adequate quality to inform primary economic modelling. An economic model was therefore developed to assess the relative cost effectiveness of interventions for the treatment of PTSD in children and young people in the UK.

### **Economic modelling methods**

### **Population**

The study population of the economic model comprised children and young people (under 18 years of age) with PTSD, who initiate treatment for PTSD in a community setting, although they may receive care in other settings over the time horizon of the analysis. This was decided because the majority of children and young people with PTSD initiate treatment for PTSD in a community setting in UK routine practice.

No distinction was made between children and young people with single trauma and those with multiple traumas as there was no evidence that the effectiveness of interventions was affected by this factor.

#### Interventions assessed

The range of interventions assessed in the economic analysis was determined by the availability of relevant clinical data included in the guideline systematic review of interventions for the treatment of PTSD in children and young people. Network meta-analysis (NMA) was employed for synthesis of the available efficacy data. Details of the NMA undertaken to inform the economic analysis are provided in the 'Efficacy data and methods of evidence synthesis' section. The guideline economic analysis assessed interventions for children and young people with PTSD that were connected to the network of evidence and were thus included in the NMA. The network included only psychological interventions, and therefore the NMA and the economic analysis explored the relative clinical and cost effectiveness of psychological interventions for the treatment of PTSD in children and young people; no pharmacological or combined interventions were assessed, as these have not been compared with psychological interventions. The NMA and the economic analysis considered separately interventions that belonged to the trauma-focused cognitive behavioural therapy (TF-CBT) class, as individual interventions had different intervention costs and, potentially, different efficacy. Based on the advice of the committee, the economic

analysis included interventions that had been tested on at least 40 individuals across RCTs included in the NMA, as this was deemed as the minimum evidence that would be adequate to support a practice recommendation. The only exception was cognitive therapy, which had been tested on only 25 children; this was included in the economic analysis as the committee was interested in the relative clinical and cost effectiveness across all interventions belonging in the TF-CBT class, and adequate evidence on TF-CBT class, which could be extrapolated to cognitive therapy, was available for other interventions within the class.

Consequently, the following psychological interventions were considered in the economic analysis of interventions for the treatment of children and young people with PTSD:

- Supportive counselling
- Group CBT (TF-CBT)
- Cohen TF-CBT / Cognitive processing therapy [Cohen/CPT] (TF-CBT)
- Cognitive therapy (TF-CBT)
- Narrative exposure (TF-CBT)
- Exposure /prolonged exposure (TF-CBT)
- Eye Movement Desensitisation Reprocessing [EMDR]
- · Family therapy
- Play therapy
- Parent training
- No treatment, reflected in waitlist or no treatment arms of RCTs included in the guideline systematic review and NMA.

### **Model structure**

A hybrid decision-analytic model consisting of a decision-tree followed by a two-state Markov model was constructed using Microsoft Office Excel 2013. The model estimated the total costs and benefits associated with provision of effective treatment options in children and young people with PTSD. The structure of the model, which aimed to simulate the course of PTSD and relevant clinical practice in the UK, was also driven by the availability of clinical data.

According to the model structure, hypothetical cohorts of children and young people with PTSD were initiated on each of the treatment options assessed, including no treatment. The duration of interventions considered in the model varied between 6 and 14 weeks. For modelling purposes relating to estimation of QALYs, the duration of a full course of treatment was assumed to be 3 months (12 weeks), without this assumption affecting resource use associated with each intervention. Following a course of treatment, children and young people in each cohort either remitted (that is, they did not meet criteria for a PTSD diagnosis) or did not remit. In the next 3 months of follow-up, those who remitted ('no PTSD') could remain in remission or relapse to a PTSD state. Conversely, those who did not remit, could remain in the PTSD state or could remit (and move to a 'no PTSD' state). The two distinct periods in the decision-tree (full course of treatment and 3-month follow-up) were informed by the results of the respective NMAs (although the 3-month follow-up period was informed by the results of the NMA only in a sensitivity analysis, as discussed later). The length of the follow-up period immediately post-treatment was set at 3 months as this was the period for which most RCT follow-up data were available across interventions.

After that point, children and young people in each cohort were entered into the Markov component of the economic model, in either the 'PTSD' or the 'no PTSD' health states,

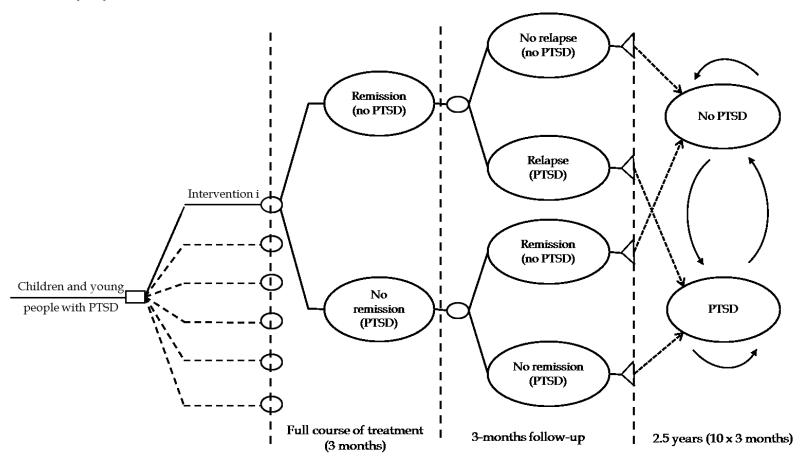
depending on their state at the end of the decision-tree. In each cycle of the Markov model, they could remain in the same health state or move between the two states of 'PTSD' and 'no PTSD'. The Markov model was run in 3-month cycles, for consistency with the duration of the two periods of the decision-tree, that is, a full course of treatment (which lasted, on average, 3 months) and another 3-month follow-up period (the length of which was determined by data availability). A half-cycle correction was applied. Due to lack of long-term comparative clinical data, transitions between the two health states in the Markov component of the model were assumed to be independent of the intervention received at the decision-tree part of the model.

The time horizon of the analysis was 3 years, consisting of the 6 months of the decision tree and another 2.5 years (10 x 3-month cycles) in the Markov component of the economic model. This time frame was considered to be long enough to capture longer-term costs and effects of treatment, without significant extrapolation over the course of PTSD.

Death was not considered in the model as there was no published evidence that mortality in children and young people with PTSD is higher than that of those in the general population. Moreover, overall mortality in children and young people is low compared with the adult population, so that the impact of a potential increase in the mortality of children and young people due to PTSD on the cost effectiveness of interventions assessed over the time horizon of the analysis was considered to be negligible.

The structure of the economic model for interventions for treatment of PTSD in children and young people is shown in Figure 172.

Figure 172. Schematic diagram of the economic model structure: interventions for the treatment of PTSD in children and young people



### Costs and outcomes considered in the analysis

The economic analysis adopted the perspective of the NHS and personal social services (PSS), as recommended by NICE (NICE, 2014). Costs consisted of intervention costs (healthcare professional time) as well as other costs incurred by children and young people with PTSD who did not remit following treatment or who experienced a relapse following remission and costs incurred by those children who were in remission, including primary, community and secondary health care and personal social services. The cost year was 2017.

The measure of outcome was the Quality Adjusted Life Year (QALY), which incorporated utilities associated with the health states of PTSD and no PTSD.

### Efficacy data and methods of evidence synthesis

### Selection of efficacy data and methods of evidence synthesis

Efficacy data for the interventions for the treatment of PTSD in children and young people that were considered in the economic modelling were derived from the respective guideline systematic review and meta-analysis of RCTs. Two types of efficacy data were extracted from the RCTs included in the review and synthesised in the guideline meta-analyses:

- Continuous data in the form of changes in PTSD symptom scores between baseline and follow-up
- Dichotomous data, either response or remission

Although the latter are more suitable for use in economic modelling as they can be directly translated into probabilities of events that correspond directly to the model health states, the remission data reported in the RCTs included in the guideline systematic review were sparse and not available for all interventions assessed in the economic analysis: continuous PTSD symptom change score data at treatment endpoint were available for 17 interventions assessed in 29 studies; on the other hand, only 9 studies reported dichotomous remission at treatment endpoint, and such data were available only for 7 interventions. Consequently, available remission data were not adequate to inform all interventions of interest included in the economic model. In contrast, continuous PTSD symptom data constituted a wider and more comprehensive evidence base that was available for a wider range of interventions. Therefore, it was decided to synthesise continuous data and to transform the analysis outputs in a suitable way, as described later, so as to inform the economic model. Two analyses of continuous data were conducted; one utilised PTSD symptom change scores between baseline and treatment endpoint and the other utilised PTSD symptom change scores between baseline and 1-4 month follow-up. Dichotomous remission data were also synthesised and utilised in a secondary economic analysis, to explore whether their consideration would alter conclusions from the base-case analysis that utilised continuous PTSD symptom change scores.

Both continuous symptom scale score data and dichotomous remission data were synthesised using network meta-analytic techniques. Network meta-analysis (NMA) is a generalisation of standard pairwise meta-analysis for A versus B trials, to data structures that include, for example, A versus B, B versus C, and A versus C trials (Dias 2011a; Lu & Ades, 2004). A basic assumption of NMA methods is that direct and indirect evidence estimate the same parameter, that is, the relative effect between A and B measured directly from an A versus B trial, is the same with the relative effect between A and B estimated indirectly from A versus C and B versus C trials. NMA techniques strengthen inference concerning the relative effect of two treatments by including both direct and indirect

comparisons between treatments, and, at the same time, allow simultaneous inference on all treatments examined in the pairwise trial comparisons while respecting randomisation (Caldwell 2005; Lu & Ades 2004). Moreover, the NMA approach assumes that the populations included in all trials are similar and thus the treatment effects are exchangeable across all populations included in the NMA (Mavridis 2015). Simultaneous estimation of the relative effects of any number of treatments is possible provided that treatments participate in a single 'network of evidence', that is, every treatment is linked to at least one of the other treatments under assessment through direct comparisons.

NMAs were conducted within a Bayesian framework using Markov Chain Monte Carlo simulation techniques implemented in WinBUGS 1.4.3 (Lunn 2000; Spiegelhalter 2003).

For the synthesis of continuous data (changes in PTSD scale score), a generalised linear model (GLM) with a normal likelihood and identity link was used (Dias et al., 2011a and 2018). Because the RCTs included in the NMAs used different continuous scales to report change in PTSD symptoms, pooling of the differences in means across different scales was not appropriate. For this reason results were expressed in the form of the Standardised Mean Difference (SMD), where the mean difference is divided by a standardising constant, which can be the population standard deviation for each scale (if known), or its estimate, often obtained by pooling the estimated standard deviations across all arms of the study (Cooper 2009). Pooling of continuous data in the NMAs utilised the Cohen's d SMD measure (Cohen 1969).

The economic model required probabilities of effect (remission). SMD cannot be directly used to estimate these probabilities. However, it was possible to transform the results of the NMAs, expressed on the SMD scale, to a log-odds ratio of effect using the following formula (Chinn 2000):

$$LOR = -\frac{\pi}{\sqrt{3}} SMD$$

This transformation assumes that remission status is determined based on a scale with an underlying normal distribution that was dichotomised into a PTSD diagnosis vs no PTSD diagnosis ('remission') using a hypothetical cut-off point on the scale.

The log-odds ratios of remission of each intervention versus no treatment (which served as the baseline treatment) were exponentiated into odds ratios. Subsequently, the probability of remission for each intervention, which was utilised in the economic model, was estimated using the following formulae:

$$intervention\ prob = \frac{odds}{(1+odds)}$$
 (1)

and

$$odds = \frac{baseline\ prob}{(1-baseline\ prob)}\ OR\ (2)$$

where baseline prob is the probability of remission for the baseline treatment (no treatment), OR is the odds ratio of remission for each intervention versus waitlist (no treatment) as estimated following exponentiation of the log-odds ratios obtained from the NMA, and odds is the odds of each intervention to achieve remission.

The WinBUGS code used to synthesise the continuous data (changes in PTSD symptom scale scores), for both random and fixed effect models, is shown in Table 35 (adapted from Dias et al., 2018). The suitability of both fixed and random effect models was assessed and compared. For the random effects model, code for both informative and uninformative prior distribution of the between-study standard deviation is provided, as the analysis on changes in PTSD symptom scores between baseline and 1-4 month follow-up suggested that the prior distribution has had some influence on the estimate of the between trial heterogeneity; therefore, an analysis utilising an informative prior distribution of the between-study standard deviation was conducted in this dataset to inform the economic analysis. The prior distribution that informed the between-study heterogeneity at the 1-4 month follow-up was derived from a study that analysed continuous mental health outcomes from meta-analyses of non-pharmacological interventions derived from the Cochrane Database of Systematic Reviews (Rhodes et al., 2015).

Table 35. WinBUGS code used to synthesise continuous data (changes in PTSD symptom scale scores) in the NMAs that informed the guideline economic modelling of interventions for the treatment of PTSD in children and young people

### Normal likelihood and identity link model

### RANDOM EFFECTS MODEL

```
# Normal likelihood, identity link: SMD with arm-based means;
# output as log Odds Ratios
# Random effects model for multi-arm trials
                            # *** PROGRAM STARTS
model{
for(i in 1:ns){
                            # LOOP THROUGH STUDIES
 w[i,1] <- 0 # adjustment for multi-arm trials is zero for control arm
 delta[i,1] <- 0
                            # treatment effect is zero for control arm
 mu[i] \sim dnorm(0,.0001)
                                  # vague priors for all trial baselines
# CONTINUOUS DATA AS ARM MEANS
for(i in 1:ns){
 # calculate pooled.sd and adjustment for SMD
 df[i] <- sum(n[i,1:na[i]]) - na[i] # denominator for pooled.var
 Pooled.var[i] <- sum(nvar[i,1:na[i]])/df[i]
 Pooled.sd[i] <- sqrt(Pooled.var[i]) # pooled sd for study i, for SMD
 H[i] <- 1
                           # use Cohen's d (ie no adjustment)
 for (k in 1:na[i]){
  se[i,k] \leftarrow sd[i,k]/sqrt(n[i,k])
  var[i,k] \leftarrow pow(se[i,k],2)
                                # calculate variances
                              # set precisions
  prec[i,k] <- 1/var[i,k]
  y[i,k] ~ dnorm(phi[i,k], prec[i,k]) # normal likelihood
  phi[i,k] <- theta[i,k] * (Pooled.sd[i]/H[i]) # theta is standardised mean
  theta[i,k] <- mu[i] + delta[i,k] # model for linear predictor, delta is SMD
  dev[i,k] \leftarrow (y[i,k]-phi[i,k])*(y[i,k]-phi[i,k])*prec[i,k]
  nvar[i,k] \leftarrow (n[i,k]-1) * pow(sd[i,k],2) # for pooled.sd
 # summed residual deviance contribution for this trial
 resdev[i] <- sum(dev[i,1:na[i]])
}
```

### Normal likelihood and identity link model

```
# RE MODEL USING UNINFORMATIVE PRIOR FOR THE BETWEEN-STUDY STANDARD
DEVIATION
                            # LOOP THROUGH ALL STUDIES
for(i in 1:ns){
 for (k in 2:na[i]){
                             # LOOP THROUGH ARMS
  # trial-specific RE distributions
  delta[i,k] ~ dnorm(md[i,k], taud[i,k])
  md[i,k] \leftarrow d[t[i,k]] - d[t[i,1]] + sw[i,k]
  # precision of RE distributions (with multi-arm trial correction)
  taud[i,k] <- tau *2*(k-1)/k
  # adjustment, multi-arm RCTs
  w[i,k] \leftarrow delta[i,k] - d[t[i,k]] + d[t[i,1]]
  # cumulative adjustment for multi-arm trials
  sw[i,k] <-sum(w[i,1:k-1])/(k-1)
 }
}
#
totresdev <- sum(resdev[])
                                   # Total Residual Deviance (all data)
# Priors distributions
d[1]<-0
                       # treatment effect is zero for control arm
# vague prior for treatment effects
for (k \text{ in } 2:nt) \{ d[k] \sim dnorm(0, .0001) \}
sdev \sim dunif(0,5)
                              # vague prior for between-trial SD
tau <- pow(sdev,-2)
                                # between-trial precision
for (c in 1:(nt-1)){
 for (k in (c+1):nt){
  diff[c,k] \leftarrow d[k] - d[c]
                             # all pairwise differences (SMD)
  lor[c,k] \leftarrow diff[c,k]^*(-3.1416/sqrt(3)) \# convert to lor (note sign)
 }
}
# RE MODEL USING INFORMATIVE PRIOR FOR THE BETWEEN-STUDY STANDARD
DEVIATION
                            # LOOP THROUGH ALL STUDIES
for(i in 1:ns){
                             # LOOP THROUGH ARMS
 for (k in 2:na[i]){
  # trial-specific RE distributions
  delta[i,k] ~ dnorm(md[i,k], taud[i,k])
  md[i,k] \leftarrow d[t[i,k]] - d[t[i,1]] + sw[i,k]
  # precision of RE distributions (with multi-arm trial correction)
  taud[i,k] <- invtausq *2*(k-1)/k
  # adjustment, multi-arm RCTs
  w[i,k] \leftarrow d[t[i,k]] + d[t[i,1]]
  # cumulative adjustment for multi-arm trials
  sw[i,k] <-sum(w[i,1:k-1])/(k-1)
 }
}
totresdev <- sum(resdev[])
                                   # Total Residual Deviance (all data)
# Priors distributions
```

```
Normal likelihood and identity link model
                    # treatment effect is zero for control arm
d[1]<-0
# vague prior for treatment effects
for (k \text{ in } 2:nt) \{ d[k] \sim dnorm(0, .0001) \}
#informative prior for log(tau-squared)
invtausq <- 1/tausq
                                       #between-study precision
                              #between-study variance
tausq <- exp(log.tausq)
sdev <- pow(tausq,0.5)
                        #between-study standard deviation
prior.prec <- pow(1.93,-2)
                                      #precision of prior distribution
#informative prior on log-between-study variance (t(-3.85,1.93^2,5))
log.tausq ~ dt(-3.85,prior.prec,5)
for (c in 1:(nt-1)){
 for (k in (c+1):nt){
                          # all pairwise differences (SMD)
  diff[c,k] \leftarrow d[k] - d[c]
  lor[c,k] \leftarrow diff[c,k]^*(-3.1416/sqrt(3)) \# convert to lor (note sign)
               or[c,k] \leftarrow exp(lor[c,k])
}
}
# rank treatments
for (k in 1:nt) {
 rk[k] <- rank(d[],k)
 best[k] <- equals(rk[k],1) # Smallest is best (i.e. rank 1)
 # prob treat k is h-th best, prob[1,k]=best[k]
 for (h in 1:nt) { prob[h,k] \leftarrow equals(rk[k],h) }
}
                     # *** PROGRAM ENDS
}
Initial values for each chain
- changes in PTSD symptom scale scores between baseline and treatment endpoint
# chain 1
mu = c(0,0,0,0,0, 0, 0,0,0,0,0, 0,0,0,0,0, 0,0,0,0,0, 0,0,0,0,0, 0,0,0,0),
sdev = 1)
# chain 2
list(d = c(NA,-1,1,1,-0.5, 1,1,1,-1,-0.7, 1,-1,0.5,0.7,-1, -1,0.5)
0.6, -0.5, -0.6, 1),
sdev = 1)
- changes in PTSD symptom scale scores between baseline and 1-4-month follow-up
[uninformative prior for the between-study standard deviation]
# chain 1
mu = c(0,0,0,0,0,0,0,0,0,0,0)
sdev = 1)
# chain 2
list(d = c(NA,-1,1,1,-0.5, 1,1,1,-1,-0.7, -1,0.5),
mu = c(0.5,1,0.7,1,-1, -0.5,0,1,-0.5,-1),
sdev = 1)
```

#### Normal likelihood and identity link model

- changes in PTSD symptom scale scores between baseline and 1-4-month follow-up [informative prior for the between-study standard deviation]

```
FIXED EFFECTS MODEL
# Normal likelihood, identity link: SMD with arm-based means;
# output as log Odds Ratios
# Fixed effect model
                            # *** PROGRAM STARTS
model{
                             # LOOP THROUGH STUDIES
for(i in 1:ns){
                                   # vague priors for all trial baselines
 mu[i] \sim dnorm(0,.0001)
# CONTINUOUS DATA AS ARM MEANS
 # calculate pooled.sd and adjustment for SMD
 df[i] <- sum(n[i,1:na[i]]) - na[i] # denominator for pooled.var
 Pooled.var[i] <- sum(nvar[i,1:na[i]])/df[i]
 Pooled.sd[i] <- sqrt(Pooled.var[i]) # pooled sd for study i, for SMD
# H[i] <- 1 - 3/(4*df[i]-1)
                                # use Hedges' g
                            # use Cohen's d (ie no adjustment)
 H[i] <- 1
 for (k in 1:na[i]){
  se[i,k] \leftarrow sd[i,k]/sqrt(n[i,k])
  var[i,k] \leftarrow pow(se[i,k],2)
                                 # calculate variances
  prec[i,k] <- 1/var[i,k]
                               # set precisions
  y[i,k] ~ dnorm(phi[i,k], prec[i,k]) # normal likelihood
  phi[i,k] \mathrel{<\!\!\!\!-} theta[i,k] * (Pooled.sd[i]/H[i]) \# theta is standardised mean
  theta[i,k] <- mu[i] + d[t[i,k]] - d[t[i,1]] # model for linear predictor
  dev[i,k] \leftarrow (v[i,k]-phi[i,k])*(v[i,k]-phi[i,k])*prec[i,k]
  nvar[i,k] \leftarrow (n[i,k]-1) * pow(sd[i,k],2) # for pooled.sd
 # summed residual deviance contribution for this trial
 resdev[i] <- sum(dev[i,1:na[i]])
totresdev <- sum(resdev[])
                                     # Total Residual Deviance (all data)
# Priors distributions
                        # treatment effect is zero for control arm
d[1]<-0
# vague prior for treatment effects
for (k \text{ in } 2:nt) \{ d[k] \sim dnorm(0, .0001) \}
for (c in 1:(nt-1)){
 for (k in (c+1):nt){
                              # all pairwise differences (SMD)
  diff[c,k] \leftarrow d[k] - d[c]
  lor[c,k] \leftarrow diff[c,k]^*(-3.1416/sqrt(3)) \# convert to lor (note sign)
 }
```

```
Normal likelihood and identity link model
# rank treatments
for (k in 1:nt) {
 rk[k] <- rank(d[],k)
 best[k] <- equals(rk[k],1) # Smallest is best (i.e. rank 1)
# prob treat k is h-th best, prob[1,k]=best[k]
for (h in 1:nt) { prob[h,k] \leftarrow equals(rk[k],h) }
                  # *** PROGRAM ENDS
}
Initial values for each chain
- changes in PTSD symptom scale scores between baseline and treatment endpoint
# chain 1
mu = c(0,0,0,0,0, 0,0,0,0,0, 0,0,0,0,0, 0,0,0,0,0, 0,0,0,0,0,0))
# chain 2
list(d = c(NA,-1,1,1,-0.5, 1,1,1,-1,-0.7, 1,-1,0.5,0.7,-1, -1,0.5)
0.6, -0.5, -0.6, 1)
- changes in PTSD symptom scale scores between baseline and 1-4-month follow-up
# chain 1
mu = c(0,0,0,0,0, 0,0,0,0,0)
# chain 2
list(d = c(NA,-1,1,1,-0.5, 1,1,1,-1,-0.7, -1,0.5),
mu = c(0.5,1,0.7,1,-1, -0.5,0,1,-0.5,-1))
```

For the synthesis of dichotomous data (remission), a binomial likelihood and logit link model was used (Dias et al., 2011a). The output of this analysis was the log-odds ratios between all pairs of interventions assessed. The log-odds ratios of remission of each intervention versus no treatment (which served as the baseline treatment) were exponentiated into odds ratios and subsequently applied onto the baseline probability of remission using the formulae (1) and (2) above, in order to obtain the absolute probability of remission for each intervention, which was utilised in the economic model.

The WinBUGS code used to synthesise the dichotomous remission data, for both random and fixed effect models, is shown in Table 36 (adapted from Dias et al., 2011a). The suitability of both models was assessed and compared. The random effects model utilised uninformative prior parameters.

Table 36. WinBUGS code used to synthesise dichotomous data (remission) in the NMAs that informed the guideline economic modelling of interventions for the treatment of PTSD in children and young people

Binomial likelihood and logit link model

#### RANDOM EFFECTS MODEL

```
# Binomial likelihood, logit link
# Random effect model, multi-arm trials
                                                  # *** PROGRAM STARTS
model{
                                                   # LOOP THROUGH STUDIES
for(i in 1:ns){
 w[i,1] <- 0
                                                  # adjustment for multi-arm trials is zero for
control arm
```

```
Binomial likelihood and logit link model
                                                          # treatment effect is zero for control arm
 delta[i,1] <- 0
                                                           # vague priors for all trial baselines
 mu[i] \sim dnorm(0,.0001)
                                                          # LOOP THROUGH ARMS
 for (k in 1:na[i]) {
                                                          # binomial likelihood
   r[i,k] \sim dbin(p[i,k],n[i,k])
                                                            # model for linear predictor
   logit(p[i,k]) \leftarrow mu[i] + delta[i,k]
                                                           # expected value of the numerators
   rhat[i,k] \leftarrow p[i,k] * n[i,k]
   dev[i,k] \leftarrow 2 * (r[i,k] * (log(r[i,k])-log(rhat[i,k]))
                                                             #Deviance contribution
      + (n[i,k]-r[i,k]) * (log(n[i,k]-r[i,k]) - log(n[i,k]-rhat[i,k])))
 }
 resdev[i] <- sum(dev[i,1:na[i]])
                                                   # summed residual deviance contribution for this trial
                                                       # LOOP THROUGH ARMS
 for (k in 2:na[i]) {
   delta[i,k] ~ dnorm(md[i,k],taud[i,k])
                                                        # trial-specific LOR distributions
                                                       # mean of LOR distributions (with multi-arm
   md[i,k] \leftarrow d[t[i,k]] - d[t[i,1]] + sw[i,k]
correction)
    taud[i,k] <- tau *2*(k-1)/k
                                                       # precision of LOR distributions (with multi-arm
correction)
   w[i,k] \leftarrow (delta[i,k] - d[t[i,k]] + d[t[i,1]])
                                                       # adjustment for multi-arm RCTs
                                                        # cumulative adjustment for multi-arm trials
   sw[i,k] <- sum(w[i,1:k-1])/(k-1)
 }
                                                           #Total Residual Deviance
totresdev <- sum(resdev[])
d[1]<- 0
                                                         # treatment effect is zero for reference
treatment
for (k \text{ in } 2:nt) \{ d[k] \sim dnorm(0,.0001) \}
                                                             # vague priors for treatment effects
sd \sim dunif(0,2)
tau <- pow(sd,-2)
# pairwise ORs and LORs for all possible pair-wise comparisons
for (c in 1:(nt-1)) { for (k in (c+1):nt) {
     or[c,k] \leftarrow exp(d[k] - d[c])
     lor[c,k] \leftarrow (d[k]-d[c])
    }
}
# ranking
for (k in 1:nt) {
  rk[k] <- nt+1-rank(d[],k)
                                           # assumes events are "good"
  best[k] <- equals(rk[k],1)
                                           #calculate probability that treat k is best
}
}
                                                         # *** PROGRAM ENDS
Initial values for each chain
#chain 1
list(d=c(NA,0,0,0,0,0,0), sd=1,
mu=c(0,0,0,0,0,0,0,0,0))
#chain 2
list(d=c(NA,0.1,-1,-0.2,1,0.1,-1), sd=0.5,
mu=c(1,-1,-2,0,0,-2,1,0,2))
```

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**FIXED EFFECTS MODEL** 

```
Binomial likelihood and logit link model
# Binomial likelihood, logit link, MTC
# Fixed effect model
model{
                                                       # *** PROGRAM STARTS
for(i in 1:ns){
                                                        # LOOP THROUGH STUDIES
 mu[i] \sim dnorm(0,.0001)
                                                         # vague priors for all trial baselines
 for (k in 1:na[i]) {
                                                        # LOOP THROUGH ARMS
  r[i,k] \sim dbin(p[i,k],n[i,k])
                                                        # binomial likelihood
  logit(p[i,k]) \leftarrow mu[i] + d[t[i,k]] - d[t[i,1]]
                                                         # model for linear predictor
  rhat[i,k] \leftarrow p[i,k] * n[i,k]
                                                         # expected value of the numerators
  dev[i,k] <- 2 * (r[i,k] * (log(r[i,k])-log(rhat[i,k]))
                                                            #Deviance contribution
     + (n[i,k]-r[i,k]) * (log(n[i,k]-r[i,k]) - log(n[i,k]-rhat[i,k])))
 }
                                              # summed residual deviance contribution for this trial
 resdev[i] <- sum(dev[i,1:na[i]])
totresdev <- sum(resdev[])
                                                #Total Residual Deviance
                                             # treatment effect is zero for reference treatment
d[1]<- 0
for (k in 2:nt) { d[k] \sim dnorm(0,.0001) }
                                                 # vague priors for treatment effects
# pairwise ORs and LORs for all possible pair-wise comparisons
for (c in 1:(nt-1)) { for (k in (c+1):nt) {
    or[c,k] \leftarrow exp(d[k] - d[c])
    lor[c,k] \leftarrow (d[k]-d[c])
    }
}
# ranking
for (k in 1:nt) {
  rk[k] <- nt+1-rank(d[],k)
                                          # assumes events are "good"
  best[k] <- equals(rk[k],1)
                                          #calculate probability that treat k is best
}
                                                        # *** PROGRAM ENDS
Initial values for each chain
#chain 1
list(d=c(NA,0,0,0,0,0,0))
mu=c(0,0,0,0,0,0,0,0,0))
#chain 2
list(d=c(NA,0.1,-1,-0.2,1,0.1,1),
mu=c(1,-1,-2,0,0,-2,1,0,2))
```

Goodness of fit of each model was assessed by comparing the posterior mean of the total residual deviance (totresdev) with the number of data points in the model. Models were also compared using the deviance information criterion (DIC), a measure of model fit penalised for model complexity, where lower values are preferred (Dias et al., 2011a; Spiegelhalter et al., 2002). Details on the interventions, data and type of model used (i.e. fixed or random effects) in each NMA are reported in the respective subheadings under the 'Efficacy data and methods of evidence synthesis' section. Each WinBUGS model was run with an initial burn-in period of 100,000 iterations, followed by 300,000 further iterations, thinned by 30 so as to obtain 10,000 iterations for use in the probabilistic economic model. Two different sets

of initial values were used; convergence was assessed by visually inspecting the mixing of the two chains in the history plots and the Brooks Gelman-Rubin diagram in WinBUGS.

Consistency between indirect and direct evidence was explored statistically by comparing the fit of a model assuming consistency with a model which allowed for inconsistency (also known as an unrelated mean effects model). The latter is equivalent to having separate, unrelated meta-analyses for every pair-wise contrast but assumes a common between-study heterogeneity across all comparisons. If the inconsistency model had a meaningfully smaller posterior mean residual deviance or heterogeneity then this indicated potential inconsistency in the data. Deviance plots, in which the posterior mean deviance of the individual data points in the inconsistency model were plotted against their posterior mean deviance in the consistency model, were inspected in order to identify studies which may have contributed to loops of evidence where inconsistency may be present. Further checks were conducted using a node-split approach implemented in R using the *gemtc* package in R (Dias et al., 2011b; van Valkenhoef & Kuiper, 2016).

When evidence of inconsistency was found, studies contributing to loops of evidence where there might be inconsistency were checked for data accuracy and analyses were repeated if corrections in the data extraction were made. However, if evidence of inconsistency was still present following any data corrections, no studies were excluded from the analysis, as their results could not be considered as less valid than those of other studies solely because of the inconsistency findings. Nevertheless, the presence of inconsistency in the NMA was highlighted and results were interpreted accordingly by the committee.

A critique of the NMA models by the NICE Technical Support Unit (TSU) including details of the inconsistency checks undertaken is provided in Appendix M.

# Synthesis of changes in PTSD symptom scores between baseline and treatment endpoint

The NMA of changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD included 29 studies and 17 interventions. Prioritisation of clinical scales for inclusion in the analysis followed the prioritisation of scales considered in the guideline systematic review and pairwise meta-analysis. Intention-to-treat (ITT) data, obtained after imputation of missing data, were prioritised over completers' data, if both were available in the same study, in accordance with the guideline systematic review protocols. For the NMA, self-reported scales were prioritised over clinician-rates scales if both were available in the same study, following advice from the committee.

Table 37 provides all studies and data considered in the NMA of changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD, while Figure 173 shows the respective network of interventions. Table 38 shows the interventions with their NMA codes, the numbers of participants randomised to each intervention across all trials included in the NMA, and the number of studies that tested each intervention.

Table 37: RCTs, interventions and efficacy data considered in the NMA of changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD

| Study        | t1 | y1    | sd1   | n1 | t2 | у2    | sd2   | n2 | t3 | у3     | sd3   | n3 |
|--------------|----|-------|-------|----|----|-------|-------|----|----|--------|-------|----|
| Chen 2014    | 1  | 1.55  | 9.01  | 12 | 3  | -2.80 | 8.37  | 10 | 5  | -14.00 | 19.94 | 10 |
| de Roos 2017 | 1  | -6.02 | 15.82 | 18 | 8  | -34.3 | 16.22 | 42 | 10 | -32.24 | 14.20 | 43 |
| Jaycox 2009  | 1  | -1.09 | 7.63  | 37 | 5  | -3.74 | 6.89  | 39 | NA | NA     | NA    | NA |

| Study   | t1 | y1     | sd1   | n1 | t2 | y2     | sd2   | n2 | t3 | у3    | sd3  | n3 |
|---|----|--------|-------|----|----|--------|-------|----|----|-------|------|----|
| Meiser-<br>Stedman 2010<br>/2017                  | 1  | -5.8   | 10.59 | 13 | 7  | -24.9  | 6.95  | 13 | NA | NA    | NA   | NA |
| Pityaratstian<br>2015                             | 1  | 0.39   | 9.78  | 18 | 5  | -1.94  | 9.40  | 18 | NA | NA    | NA   | NA |
| Smith 2007  | 1  | -6.3   | 9.63  | 11 | 7  | -39    | 7.65  | 12 | NA | NA    | NA   | NA |
| Auslander<br>2016                                 | 2  | 0.8    | 9.68  | 10 | 5  | -5.68  | 6.71  | 15 | NA | NA    | NA   | NA |
| Goldbeck<br>2016 /Sachser<br>2016                 | 1  | -7.52  | 9.18  | 82 | 6  | -13.4  | 9.63  | 74 | NA | NA    | NA   | NA |
| Jensen 2014                                       | 2  | -10.01 | 7.63  | 63 | 6  | -15.48 | 6.96  | 59 | NA | NA    | NA   | NA |
| Langley 2015                                      | 1  | -2.05  | 9.82  | 36 | 5  | -14.41 | 9.91  | 35 | NA | NA    | NA   | NA |
| Shein-Szydlo<br>2016                              | 1  | -1.94  | 9.84  | 49 | 6  | -23.72 | 8.12  | 50 | NA | NA    | NA   | NA |
| Stein 2003a<br>/Kataoka 2011                      | 1  | -8     | 7.01  | 63 | 5  | -15.6  | 5.07  | 54 | NA | NA    | NA   | NA |
| Al-Hadethe<br>2015                                | 1  | 2.1    | 7.25  | 20 | 8  | -5.05  | 5.64  | 19 | 17 | -9.95 | 5.37 | 20 |
| Deblinger<br>1996/1999                            | 2  | -3.29  | 2.34  | 14 | 9  | -5.48  | 2.12  | 21 | 15 | -4.7  | 2.34 | 20 |
| King 2000   | 1  | -1.47  | 1.68  | 12 | 9  | -5.75  | 3.01  | 12 | 16 | -7.08 | 4.10 | 12 |
| Ruf 2010  | 1  | -4.5   | 12.34 | 13 | 8  | -26.1  | 9.75  | 12 | NA | NA    | NA   | NA |
| Gilboa-<br>Schechtman<br>2004/2010                | 3  | -10.79 | 8.36  | 19 | 9  | -19.37 | 8.45  | 19 | NA | NA    | NA   | NA |
| Cohen 1998<br>/2005a                              | 3  | -0.91  | 3.97  | 41 | 6  | -1.85  | 3.56  | 41 | NA | NA    | NA   | NA |
| Cohen 2011<br>/2005b                              | 3  | -1.66  | 9.14  | 60 | 6  | -7.16  | 13.52 | 64 | NA | NA    | NA   | NA |
| Foa 2013  | 3  | -15.3  | 6.83  | 30 | 9  | -18.7  | 6.86  | 31 | NA | NA    | NA   | NA |
| Ford 2012   | 3  | -17    | 9.53  | 20 | 6  | -24.4  | 13.93 | 26 | NA | NA    | NA   | NA |
| Diehle 2015<br>/Lindauer 2009                     | 6  | -20.2  | 15.58 | 23 | 10 | -20.9  | 20.08 | 25 | NA | NA    | NA   | NA |
| Soberman<br>2002                                  | 2  | -5.73  | 12.39 | 11 | 11 | -5.5   | 10.20 | 10 | NA | NA    | NA   | NA |
| Ahmad 2007<br>/2008                               | 1  | -7.4   | 14.01 | 16 | 10 | -6.3   | 15.35 | 17 | NA | NA    | NA   | NA |
| Lieberman<br>2005 / 2006 /<br>Ghosh Ippen<br>2011 | 15 | -0.4   | 3.03  | 29 | 12 | -3.61  | 2.33  | 36 | NA | NA    | NA   | NA |
| Kazak 2004  | 1  | -4.49  | 5.53  | 74 | 13 | -6.53  | 5.36  | 75 | NA | NA    | NA   | NA |
| Deeba 2015  | 2  | 0.77   | 6.00  | 60 | 14 | -5.2   | 5.15  | 69 | NA | NA    | NA   | NA |
| Schottelkorb<br>2012                              | 6  | -2.25  | 10.04 | 12 | 14 | -3.36  | 9.40  | 14 | NA | NA    | NA   | NA |
| Gordon 2006<br>/2008                              | 1  | -0.1   | 0.26  | 39 | 4  | -0.5   | 0.21  | 38 | NA | NA    | NA   | NA |

t1, t2, t3 indicate the coded treatment in each trial arm; codes of treatments are provided in **Table 38** y1, y2, y3 indicate the mean change in effect in each trial arm sd1, sd2, sd3 indicate the standard deviation of the mean change in effect in each trial arm

| Study  | t1 | y1 | sd1 | n1 | t2 | y2 | sd2 | n2 | t3 | у3 | sd3 | n3 |
|--|----|----|-----|----|----|----|-----|----|----|----|-----|----|
| n1, n2, n3 indicate the number of participants in each trial arm |    |    |     |    |    |    |     |    |    |    |     |    |
| NA: non-applicable   |    |    |     |    |    |    |     |    |    |    |     |    |

Figure 173. Network of interventions included in the NMA of changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD

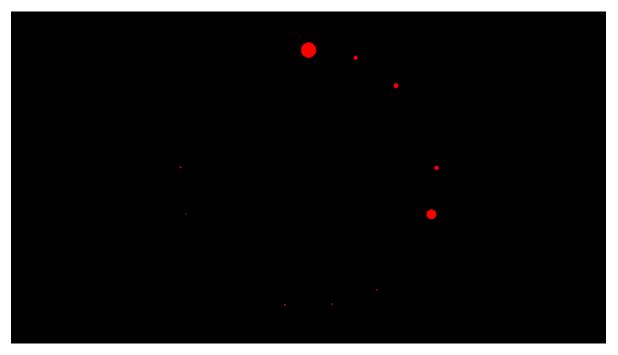


Table 38. NMA of changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD: Interventions with NMA codes, numbers of participants (N) randomised to each intervention across RCTs and number of RCTs (k) that tested each intervention

| Code | Intervention                       | N randomised<br>(N total = 1960) | k (k total = 29;<br>63 arms) |
|------|------------------------------------|----------------------------------|------------------------------|
| 1    | Waitlist / no treatment            | 513                              | 16                           |
| 2    | TAU                                | 158                              | 5                            |
| 3    | Supportive counselling             | 180                              | 6                            |
| 4    | Meditation                         | 38                               | 1                            |
| 5    | TF-CBT group CBT                   | 171                              | 6                            |
| 6    | TF-CBT Cohen/CPT                   | 349                              | 8                            |
| 7    | TF-CBT cognitive therapy           | 25                               | 2                            |
| 8    | TF-CBT narrative exposure          | 73                               | 3                            |
| 9    | TF-CBT exposure/prolonged exposure | 83                               | 4                            |
| 10   | EMDR                               | 85                               | 3                            |
| 11   | EMDR & TAU                         | 10                               | 1                            |
| 12   | Child-parent psychotherapy         | 36                               | 1                            |

| Code | Intervention                       | N randomised<br>(N total = 1960) | k (k total = 29;<br>63 arms) |
|------|------------------------------------|----------------------------------|------------------------------|
| 13   | Family therapy                     | 75                               | 1                            |
| 14   | Play therapy                       | 83                               | 2                            |
| 15   | Parent training                    | 49                               | 2                            |
| 16   | TF-CBT & parent training           | 12                               | 1                            |
| 17   | Combined somatic/cognitive therapy | 20                               | 1                            |

EMDR: eye movement desensitisation reprocessing; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

#### It is noted that:

- Waitlist and no treatment arms were included in the same node, following advice from the
  committee. However, it is acknowledged that the baseline effect of waitlist may be lower
  than that of 'no treatment' (Furukawa et al., 2014), resulting in the relative effects of active
  interventions having been potentially exaggerated in waitlist-controlled studies compared
  with their expected effects versus a 'no treatment'-control.
- Of the studies included in the guideline systematic review that reported changes in PTSD symptom scores between baseline and treatment endpoint, Lyshak-Stelzer 2007 compared art therapy + TAU vs attention placebo + TAU; Layne 2008 compared TF-CBT & psychoeducation vs psychoeducation alone. These treatments could not be connected in the network and thus were not considered in the NMA.
- The systematic review of pharmacological interventions for the treatment of PTSD in children and young people included one trial (Robb 2010) that evaluated sertraline versus pill placebo. In addition, Cohen 2007 evaluated sertraline plus TF-CBT vs placebo plus TF-CBT. Both these studies were not connected in the network and thus the respective interventions were not considered in the NMA.

# Results of the network meta-analysis: changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD

The random effects model demonstrated a better fit for the data (totresdev = 63.01; DIC = 275.27) than the fixed effects model (totresdev = 142.20; DIC = 340.17). The number of data points (study arms) in the model was 63, suggesting a good fit of the random effects model. The between-study heterogeneity was large compared with the size of treatment effects (sd 0.58). No evidence of inconsistency was identified in the network using either global inconsistency checks or the node-splitting method. Details of the inconsistency checks are provided in Appendix M.

The results of the random effects model are shown in Table 39. Interventions have been ordered from best to worst, according to their mean ranking across model iterations. Relative effects versus waitlist / no treatment (mean SMD and log-odds ratio and 95% credible intervals [CrI]) are reported. Posterior mean ranks of each intervention (and 95% CrI) are also provided, where a rank of 1 is best.

Table 39. Results of the NMA: changes in PTSD symptom scores between baseline and treatment endpoint in children and young people with PTSD (random effects model)

| Intervention                          | Mean SMD (95%<br>Crl) vs waitlist | Mean LOR (95%<br>Crl) vs waitlist | Mean ranking<br>(95% Crl) |
|---------------------------------------|-----------------------------------|-----------------------------------|---------------------------|
| TF-CBT (cognitive therapy)            | -2.94 (-3.94 to -1.95)            | 5.33 (3.54 to 7.15)               | 1.58 (1 to 4)             |
| Combined somatic/cognitive therapy    | -2.14 (-3.34 to -0.92)            | 3.88 (1.66 to 6.05)               | 3.77 (1 to 11)            |
| Child-parent psychotherapy            | -2.16 (-4.02 to -0.26)            | 3.92 (0.48 to 7.29)               | 4.13 (1 to 13)            |
| TF-CBT & parent training              | -1.79 (-3.15 to -0.45)            | 3.24 (0.81 to 5.71)               | 5.40 (1 to 14)            |
| Meditation                            | -1.67 (-2.94 to -0.41)            | 3.03 (0.75 to 5.32)               | 5.96 (1 to 14)            |
| TF-CBT (narrative exposure)           | -1.49 (-2.25 to -0.74)            | 2.71 (1.34 to 4.07)               | 6.57 (3 to 12)            |
| TF-CBT (exposure/ prolonged exposure) | -1.34 (-2.15 to -0.51)            | 2.42 (0.93 to 3.90)               | 7.51 (3 to 12)            |
| Play therapy                          | -1.35 (-2.48 to -0.20)            | 2.44 (0.37 to 4.49)               | 7.60 (2 to 14)            |
| TF-CBT (Cohen/CPT)                    | -1.17 (-1.78 to -0.54)            | 2.12 (0.99 to 3.23)               | 8.69 (5 to 13)            |
| EMDR                                  | -0.99 (-1.76 to -0.23)            | 1.80 (0.41 to 3.19)               | 10.14 (5 to 15)           |
| Parent training                       | -0.96 (-2.32 to 0.41)             | 1.74 (-0.74 to 4.21)              | 10.28 (3 to 17)           |
| TF-CBT (group CBT)                    | -0.91 (-1.48 to -0.34)            | 1.65 (0.61 to 2.68)               | 10.72 (6 to 15)           |
| Supportive counselling                | -0.59 (-1.29 to 0.12)             | 1.08 (-0.19 to 2.34)              | 12.96 (9 to 16)           |
| Family therapy                        | -0.37 (-1.60 to 0.84)             | 0.67 (-1.52 to 2.91)              | 13.59 (5 to 17)           |
| EMDR & TAU                            | -0.28 (-1.96 to 1.40)             | 0.52 (-2.54 to 3.56)              | 13.65 (4 to 17)           |
| TAU                                   | -0.31 (-1.16 to 0.56)             | 0.55 (-1.01 to 2.10)              | 14.51 (10 to 17)          |
| Waitlist / no treatment               | Reference                         | Reference                         | 15.96 (14 to 17)          |

Standard deviation: mean 0.58 (95% Crl 0.37 to 0.89)

Total residual deviance 62.93 (95% Crl 42.95 to 86.24)

Crl: credible intervals; EMDR: eye movement desensitisation reprocessing; LOR: log-odds ratio; SMD: standardised mean difference; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

Negative values for the SMD and positive values for the LOR indicate a better effect for the intervention compared with the reference treatment (waitlist / no treatment).

Interventions in italics were not considered in the economic analysis due to the low number of people randomised to each of them or due to their being interventions of no interest

Detailed results of all pair-wise comparisons between interventions are shown in Appendix N.

As reported earlier, the economic analysis included interventions that had been tested on at least 40 individuals across RCTs included in the NMA, as this was deemed as the minimum evidence that would be adequate to support a practice recommendation. The only exception was cognitive therapy, which had been tested on only 25 children; this was included in the economic analysis as the committee was interested in the relative clinical and cost effectiveness across all interventions belonging to the TF-CBT class. Therefore, meditation, child-parent psychotherapy, TF-CBT & parent training, and combined somatic/cognitive therapy were not considered in the economic analysis. Moreover, TAU and EMDR & TAU were of no interest and were thus not included in the economic analysis, as TAU was not accurately defined across the studies and varied between "access to outpatient mental"

health services" to "therapists being asked to provide the treatment they believed would be effective for the particular case with all participants receiving individual treatment".

The output of the NMA used in the economic analysis was the log-odds ratio of every intervention versus waitlist / no treatment.

# Synthesis of changes in PTSD symptom scores between baseline and 1-4 month follow-up

The NMA of changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD included 10 studies and 12 interventions. As with treatment endpoint continuous data, prioritisation of clinical scales for inclusion in the analysis followed the prioritisation of scales considered in the guideline systematic review and pairwise meta-analysis. Intention-to-treat (ITT) data, obtained after imputation of missing data, were prioritised over completers' data, if both were available in the same study, in accordance with the guideline systematic review protocols. For the NMA, self-reported scales were prioritised over clinician-rates scales if both were available in the same study, following advice from the committee.

Table 40 provides all studies and data considered in the NMA of changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD, whereas Figure 174 shows the respective network of interventions. Table 41 shows the interventions with their NMA codes, the numbers of participants randomised to each intervention across all trials included in the NMA, and the number of studies that tested each intervention.

Table 40: RCTs, interventions and efficacy data considered in the NMA of changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD

| Study                      | t1 | y1     | sd1   | n1 | t2 | y2     | sd2   | n2 | t3 | у3    | sd3   | n3 |
|----------------------------|----|--------|-------|----|----|--------|-------|----|----|-------|-------|----|
| Ahrens 2002                | 1  | 0.08   | 5.76  | 19 | 5  | -12.11 | 8.05  | 19 | NA | NA    | NA    | NA |
| Berger 2009                | 1  | -1.52  | 5.20  | 82 | 3  | -8.73  | 5.82  | 84 | NA | NA    | NA    | NA |
| Chen 2014                  | 1  | -2.2   | 9.07  | 12 | 3  | -6.5   | 10.84 | 10 | 4  | -22.8 | 8.75  | 10 |
| Pityaratstian<br>2015      | 1  | 0.78   | 10.15 | 18 | 4  | -5.67  | 8.50  | 18 | NA | NA    | NA    | NA |
| Al-Hadethe<br>2015         | 1  | 3.5    | 7.41  | 20 | 6  | -4     | 7.72  | 19 | 12 | -9.4  | 5.35  | 20 |
| Deblinger<br>1996/1999     | 2  | -4.15  | 2.90  | 14 | 7  | -5.53  | 2.09  | 21 | 10 | -5.8  | 2.29  | 20 |
| Ertl 2011 /<br>Neuner 2007 | 1  | -10.68 | 13.80 | 28 | 3  | -16.87 | 14.42 | 24 | 6  | -20.3 | 12.73 | 26 |
| King 2000                  | 1  | -1.91  | 1.95  | 12 | 7  | -4.66  | 2.52  | 12 | 11 | -6.33 | 4.06  | 12 |
| de Roos<br>2017            | 6  | -36.63 | 15.83 | 42 | 8  | -31.31 | 14.61 | 43 | NA | NA    | NA    | NA |
| Soberman<br>2002           | 2  | -6.78  | 8.14  | 11 | 9  | -12.83 | 8.1   | 12 | NA | NA    | NA    | NA |

t1, t2, t3 indicate the coded treatment in each trial arm; codes of treatments are provided in Table 41 y1, y2, y3 indicate the mean change in effect in each trial arm

NA: non-applicable

sd1, sd2, sd3 indicate the standard deviation of the mean change in effect in each trial arm

n1, n2, n3 indicate the number of participants in each trial arm

Figure 174. Network of interventions included in the NMA of changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD

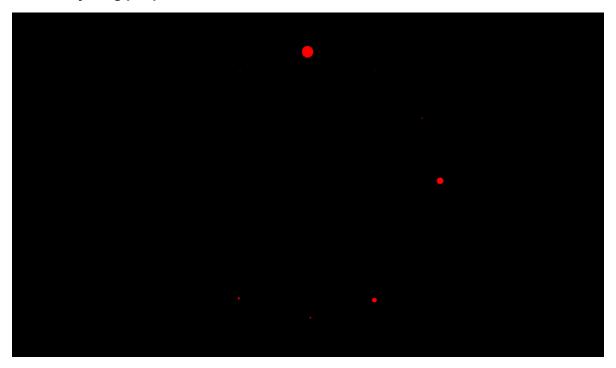


Table 41. NMA of changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD: Interventions with NMA codes, numbers of participants (N) randomised to each intervention across RCTs and number of RCTs (k) that tested each intervention

| Code | Intervention                         | N randomised<br>(N total = 608) | k (k total = 10;<br>25 arms) |
|------|--------------------------------------|---------------------------------|------------------------------|
| 1    | Waitlist / no treatment              | 191                             | 7                            |
| 2    | TAU                                  | 25                              | 2                            |
| 3    | Supportive counselling               | 34                              | 2                            |
| 4    | TF-CBT (group CBT)                   | 112                             | 3                            |
| 5    | TF-CBT (Cohen/CPT)                   | 19                              | 1                            |
| 6    | TF-CBT (narrative exposure)          | 87                              | 3                            |
| 7    | TF-CBT (exposure/prolonged exposure) | 33                              | 2                            |
| 8    | EMDR                                 | 43                              | 1                            |
| 9    | EMDR & TAU                           | 12                              | 1                            |
| 10   | Parent training                      | 20                              | 1                            |
| 11   | TF-CBT & parent training             | 12                              | 1                            |
| 12   | Combined somatic/cognitive therapy   | 20                              | 1                            |

EMDR: eye movement desensitisation reprocessing; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

# Results of the network meta-analysis: changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD

The random effects model demonstrated a better fit for the data (totresdev = 25.22; DIC = 115.57) than the fixed effects model (totresdev = 41.51; DIC = 128.18). The number of data points (study arms) in the model was 25, suggesting satisfactory fit of the random effects model. Because the distribution of the posterior between-study standard deviation suggested that the uninformative prior distribution (Uniform(0,5)) has had some influence on the estimate of heterogeneity and resulted in high heterogeneity (0.97) (Appendix M), an analysis with an informative prior distribution (Rhodes et al., 2015) was conducted to assess whether this would have an impact on the final results. This analysis (using informative priors) was used to populate the economic model.

The between-study heterogeneity, after use of informative priors, was moderate to large compared with the size of treatment effects (sd 0.51). Inconsistency checks did not indicate evidence of inconsistency.

The results of the random effects model are shown in Table 42. Interventions have been ordered from best to worst, according to their mean ranking across model iterations. Relative effects versus waitlist / no treatment (mean SMD and log-odds ratio and 95% Crl) are reported. Posterior mean ranks of each intervention (and 95% Crl) are also provided, where a rank of 1 is best.

Table 42. Results of the NMA: changes in PTSD symptom scores between baseline and 1-4 month follow-up in children and young people with PTSD (random effects model)

| Intervention                          | Mean SMD (95%<br>Crl) vs waitlist | Mean LOR (95%<br>Crl) vs waitlist | Mean ranking<br>(95% Crl) |
|---------------------------------------|-----------------------------------|-----------------------------------|---------------------------|
| Combined somatic/cognitive therapy    | -1.80 (-3.01 to -0.58)            | 3.26 (1.05 to 5.46)               | 3.02 (1 to 9)             |
| TF-CBT (Cohen/CPT)                    | -1.74 (-3.09 to -0.42)            | 3.16 (0.76 to 5.60)               | 3.37 (1 to 10)            |
| TF-CBT (group CBT)                    | -1.51 (-2.48 to -0.61)            | 2.75 (1.11 to 4.51)               | 4.09 (1 to 9)             |
| TF-CBT & parent training              | -1.49 (-2.90 to -0.07)            | 2.70 (0.13 to 5.27)               | 4.33 (1 to 10)            |
| EMDR & TAU                            | -1.10 (-3.51 to 1.23)             | 1.99 (-2.23 to 6.37)              | 6.06 (1 to 12)            |
| Parent training                       | -1.04 (-2.91 to 0.80)             | 1.89 (-1.44 to 5.28)              | 6.30 (1 to 11)            |
| TF-CBT (narrative exposure)           | -0.94 (-1.84 to -0.04)            | 1.71 (0.07 to 3.33)               | 6.85 (3 to 11)            |
| TF-CBT (exposure /prolonged exposure) | -0.92 (-2.25 to 0.37)             | 1.68 (-0.67 to 4.08)              | 6.97 (3 to 11)            |
| Supportive counselling                | -0.74 (-1.41 to 0.06)             | 1.34 (-0.10 to 2.56)              | 7.94 (4 to 11)            |
| EMDR                                  | -0.59 (-2.12 to 0.97)             | 1.08 (-1.75 to 3.85)              | 8.48 (2 to 12)            |
| TAU                                   | -0.35 (-2.26 to 1.60)             | 0.63 (-2.83 to 4.11)              | 9.52 (3 to 12)            |
| Waitlist / no treatment               | Reference                         | Reference                         | 11.08 (8 to 12)           |

Standard deviation: mean 0.51 (95% Crl 0.12 to 1.20)

Total residual deviance 26.85 (95% Crl 13.99 to 43.95)

CPT: cognitive processing therapy; CrI: credible intervals; EMDR: eye movement desensitisation reprocessing; LOR: log-odds ratio; SMD: standardised mean difference; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

Negative values for the SMD and positive values for the LOR indicate a better effect for the intervention compared with the reference treatment (waitlist / no treatment).

| Intervention  | Mean SMD (95%    | Mean LOR (95%    | Mean ranking |  |  |  |  |  |
|---|------------------|------------------|--------------|--|--|--|--|--|
|   | Crl) vs waitlist | Crl) vs waitlist | (95% Crl)    |  |  |  |  |  |
| Interventions in italics were not considered in the economic analysis |                  |                  |              |  |  |  |  |  |

Detailed results of all pair-wise comparisons between interventions are provided in Appendix N.

The committee noted the uncertainty characterising this analysis, due to the small number of studies and participants and the moderate between-study heterogeneity after use of informative priors. Therefore, the 1-4 month follow-up data (log-odds ratios of every intervention versus waitlist / no treatment) were used only in a sensitivity analysis, to obtain probabilities of remission for all active interventions during 3-6 months from treatment initiation. Follow-up data were not available for TF-CBT (cognitive therapy), play therapy and family therapy. The follow-up effect of TF-CBT (cognitive therapy) was borrowed from TF-CBT (Cohen/CPT); the follow-up effect of play and family therapy was assumed to equal that of no treatment.

In the base-case analysis the model assumed that at 3-6 months the probability of remission of each active intervention was equal to the baseline probability of remission for no treatment.

### Synthesis of dichotomous remission data at treatment endpoint

The NMA of dichotomous remission data at treatment endpoint in children and young people with PTSD included 9 studies and 7 interventions. In all studies remission was defined as loss of PTSD diagnosis according to ICD, DSM or similar criteria. Table 43 provides all studies and data considered in the NMA of dichotomous remission data at treatment endpoint in children and young people with PTSD, whereas Figure 175 shows the respective network of interventions. Table 44 shows the interventions with their NMA codes, the numbers of participants randomised to each intervention across all trials included in the NMA, and the number of studies that tested each intervention.

Table 43: RCTs, interventions and efficacy data considered in the NMA of dichotomous remission data at treatment endpoint in children and young people with PTSD

| Study                       | t1 | r1 | n1 | t2 | r2 | n2 |
|-----------------------------|----|----|----|----|----|----|
| Meiser-Stedman 2010/2017    | 1  | 3  | 15 | 4  | 10 | 14 |
| Smith 2007                  | 1  | 5  | 12 | 4  | 11 | 12 |
| Goldbeck 2016/Sachser 2016  | 1  | 24 | 63 | 5  | 34 | 57 |
| Jensen 2014                 | 2  | 23 | 42 | 5  | 28 | 36 |
| Ruf 2010                    | 1  | 4  | 13 | 6  | 11 | 13 |
| Gilboa-Schechtman 2004/2010 | 3  | 7  | 19 | 7  | 13 | 19 |
| Cohen 2011/2005b            | 3  | 8  | 18 | 5  | 24 | 32 |
| Foa 2013                    | 3  | 13 | 30 | 7  | 24 | 31 |
| Ford 2012                   | 3  | 7  | 26 | 5  | 10 | 33 |

t1, t2, t3 indicate the coded treatment in each trial arm; codes of treatments are provided in Table 44

r1, r2, r3 indicate the number of events in each trial arm

n1, n2, n3 indicate the number of participants in each trial arm

Figure 175. Network of interventions included in the NMA of dichotomous remission data at treatment endpoint in children and young people with PTSD

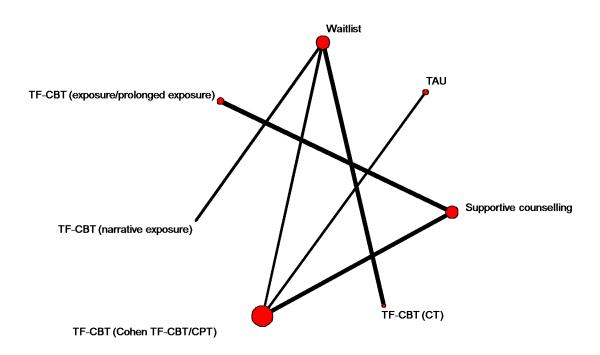


Table 44. NMA of dichotomous remission data at treatment endpoint in treatment and young people with PTSD: Interventions with NMA codes, and numbers of participants (N) randomised to each intervention across RCTs and number of RCTs (k) that tested each intervention

| Code | Intervention                         | N randomised<br>(N total = 485) | k (k total = 9;<br>18 arms) |
|------|--------------------------------------|---------------------------------|-----------------------------|
| 1    | Waitlist                             | 103                             | 4                           |
| 2    | TAU                                  | 42                              | 1                           |
| 3    | Supportive counselling               | 93                              | 4                           |
| 4    | TF-CBT (cognitive therapy)           | 26                              | 2                           |
| 5    | TF-CBT (Cohen/CPT)                   | 158                             | 4                           |
| 6    | TF-CBT (narrative exposure)          | 13                              | 1                           |
| 7    | TF-CBT (exposure/prolonged exposure) | 50                              | 2                           |

CPT: cognitive processing therapy; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

# Results of the network meta-analysis: remission at treatment endpoint in children and young people with PTSD

The fixed effects model demonstrated a good fit for the data (totresdev = 17.37; DIC = 93.71) that was comparable to the fit of the random effects model (totresdev = 17.38; DIC = 95.03). The number of data points (study arms) in the model was 18. Thus, the fixed effects model was the preferred model. Since there were no closed loops of direct evidence within the network, inconsistency checks were not possible to perform for this outcome.

The results of the fixed effects model are shown in Table 45. Interventions have been ordered from best to worst, according to their mean ranking across model iterations. Relative

effects versus waitlist / no treatment (mean log-odds ratio and 95% CrI) are reported. Posterior mean ranks of each intervention (and 95% CrI) are also provided, where a rank of 1 is best.

Table 45. Results of the NMA: dichotomous remission at treatment endpoint in children and young people with PTSD (fixed effects model)

| Intervention                         | Mean LOR (95%<br>Crl) vs waitlist | Mean ranking<br>(95% Crl) |
|--------------------------------------|-----------------------------------|---------------------------|
| TF-CBT (narrative exposure)          | 2.81 (0.87 to 5.13)               | 1.69 (1 to 4)             |
| TF-CBT (CT)                          | 2.66 (1.28 to 4.22)               | 1.72 (1 to 3)             |
| TF-CBT (exposure/prolonged exposure) | 1.62 (0.22 to 3.04)               | 2.81 (1 to 4)             |
| TF-CBT (Cohen/CPT)                   | 0.89 (0.15 to 1.64)               | 3.90 (3 to 5)             |
| Supportive counselling               | 0.15 (-0.98 to 1.28)              | 5.64 (4 to 7)             |
| Waitlist / no treatment              | Reference                         | 5.95 (5 to 7)             |
| TAU                                  | -0.21 (-1.48 to 1.03)             | 6.30 (5 to 7)             |

Total residual deviance 17.38 (95% Crl 8.51 to 30.01)

CPT: cognitive processing therapy; CrI: credible intervals; LOR: log-odds ratio; TAU: treatment as usual; TF-CBT: trauma-focused cognitive behavioural therapy

Positive values for the LOR indicate a better effect for the intervention compared with the reference treatment (waitlist / no treatment).

Interventions in italics were not considered in the economic analysis

Detailed results of all pair-wise comparisons between interventions are shown in Appendix N.

The committee noted the very small numbers randomised to most interventions, in particular cognitive therapy (N=26) and narrative exposure (N=13) and uncertainty characterising this analysis. The results of this analysis, as reported earlier, were used only in a secondary economic analysis, which aimed to explore whether the conclusions of the economic analysis based on use of dichotomous remission data would be different from those of the base-case analysis that utilised continuous PTSD symptom change scores.

#### Dichotomous remission data at 1-4 month follow-up

Dichotomous remission data at 3-month follow-up were very limited and comprised evidence on group CBT from 2 studies (Berger 2009 and Pityaratstian 2015) which compared group CBT versus waitlist. Pair-wise meta-analysis of these two studies indicated a high mean odds ratio of group CBT versus waitlist of 25.86 (95% CrI 7.28 to 91.84). This value was used in sensitivity analysis, applied onto the baseline probability of remission for no treatment at 3-6 months of the economic model, to obtain probabilities of remission for all active interventions during this period (so that all interventions were assumed to have the same effect at the 3-month follow-up due to lack of differential data); in the main secondary analysis of dichotomous remission data, the model assumed that at 3-6 months the probability of remission of each active intervention was equal to the baseline probability of remission for no treatment.

#### Other clinical input parameters

Other clinical input parameters included

- the baseline (no treatment) probability of remission, which was applied as the baseline in the decision-tree and also across all treatment options in the Markov part of the model
- the probability of relapse following remission, which was independent of the intervention received at the start of the model.

#### Baseline probability of remission in children and young people with PTSD

A number of studies were identified in the literature that reported the probability of remission over time in children and young people with PTSD (De Young 2012; Hiller 2016; Hong 2014; Kronenberg 2010; La Greca 2013a & 2013b; Rosellini 2017; Yule 2000). The majority of studies were relatively small prospective cohort studies (N ranging from 116 to 568) that explored the development and course of PTSD in children and young people following exposure to trauma over a period that varied from 6 months to 8 years (De Young et al., Hong 2014; Kronenberg 2010; La Greca 2013a & 2013b; Yule 2013). One study (Hiller 2016) was a systematic review of 27 longitudinal studies of PTSD in children and young people that assessed changes in PTSD prevalence and symptoms in this population over the first 12 months post-trauma. Finally, one study reported data on the course of PTSD derived from 1575 people with lifetime PTSD who had participated in 22 WHO World Mental Health surveys (Rosellini 2017). The study reported rates of remission of PTSD over 120 months (10 years) following PTSD onset, which was the longest follow-up period for which a sufficient number of cases were observed for stable estimation of conditional probability of remission. The publication included a graph that illustrated the probability of PTSD recovery over time for different age groups, including data on children aged 0-12 years and data on young people aged 13-24 years. The PTSD remission data from this study, as provided in the graph, were selected to populate the economic model because they were directly relevant to the study population, they were derived from a large study sample and they were available for a long time period that fully covered the time horizon of the economic analysis.

Digital software (http://www.digitizeit.de) was used to read and extract the cumulative proportions of children aged 0-12 years and young people aged 13-24 years that remitted from PTSD at 3 months, 6 months, 12 months, 24 months, and 36 months from PTSD onset. The values at each time point were averaged between the two groups, to cover the whole range of the economic analysis study population. The extracted values were used to estimate the probability of remission between 0-3 months, 3-6 months, 6-12 months, 12-24 months and 24-36 months, conditional on not having achieved remission prior to the beginning of each interval. The probability of remission between 3-6 months was very similar to that between 6-12 months, and therefore a probability of remission between 3-12 months was estimated. The estimated probabilities of remission during these time periods were subsequently transformed into 3-month probabilities that were used to inform the economic model.

Table 46 shows the estimated cumulative probability of remission for children and young people at 3, 12, 24 and 36 months from PTSD onset, the probability of remission between 0-3, 3-12, 12-24 and 24-36 months (conditional on not having achieved remission prior to the beginning of the interval), and the 3-monthly probability of remission during these time periods.

Table 46: Probability of remission overtime in children and young people with PTSD, as estimated based on data extracted from Rosellini and colleagues (2017)

| Time from PTSD onset  | Cumulative probability of remission | Time interval | Probability of remission over the time interval* | 3-monthly probability during the time interval* |  |  |
|---|-------------------------------------|---------------|--|---|--|--|
| 3 months  | 0.174                               | 0-3 months    | 0.174  | 0.174   |  |  |
| 12 months   | 0.370                               | 3-12 months   | 0.238  | 0.087   |  |  |
| 24 months   | 0.445                               | 12-24 months  | 0.118  | 0.031   |  |  |
| 36 months   | 0.500                               | 24-36 months  | 0.100  | 0.026   |  |  |
| * conditional on not having achieved remission prior to the beginning of the interval |                                     |               |  |   |  |  |

It needs to be noted that the economic analysis evaluated interventions for the delayed (>3 months) treatment of PTSD in children and young people. The economic model is thus assumed to start at month 3 from PTSD onset. The data reported in Table 46 refer to time periods from PTSD onset, meaning that the remission data corresponding to 0-3 months after PTSD onset refer to a time period just before treatment was received by the model's study population. Therefore these data were not utilised in the economic analysis. The economic model was informed by the following available data:

- The 3-month probability of remission over 3-12 months from PTSD onset informed months 0-9 of the economic model: these data were used to populate the no treatment arm during the first 6 months of the economic model, comprising 3 months of a full course of treatment plus the 3-month follow-up, i.e. over the duration of the decision-tree (months 0-6 of the economic model). It also informed all model arms in months 3-6 of the economic model in the base-case analysis. Finally, it informed all model arms in the first cycle of the Markov model (months 6-9 of the economic model), as the course of PTSD after 6 months of treatment was assumed to be independent of the treatment received.
- The 3-month probability of remission over 12-24 months from PTSD onset informed all model arms in the next 4 cycles of the Markov model (months 9-21 of the economic model).
- The 3-month probability of remission over 24-36 months from PTSD onset informed all model arms in the next 5 cycles of the Markov model (months 21-36 of the economic model); the 3-month probability was extrapolated to the period of 36-39 months from PTSD onset (i.e. months 33-36 of the economic model) for simplification.

#### Risk of relapse following remission of PTSD

No published evidence on the risk of relapse following remission from PTSD in children and young people was identified in the published literature. The committee advised that this was very low. Therefore, an annual risk of relapse of 0.10 was assumed. This was translated into a 3-month probability of relapse of 0.026, which was applied in the 3-month follow-up period of the decision-tree and over the whole duration of the Markov model. In deterministic sensitivity analysis the annual risk value of 0.10 was varied between zero and 0.20.

#### Utility data and estimation of quality adjusted life years (QALYs)

In order to express outcomes in the form of QALYs, the health states of the economic model (remission, response not reaching remission, no response or relapse) need to be linked to appropriate utility scores. Utility scores represent the HRQoL associated with specific health states on a scale from 0 (death) to 1 (perfect health); they are estimated using preference-

based measures that capture people's preferences on the HRQoL experienced in the health states under consideration.

The systematic review of utility data on PTSD-related heath states identified 2 studies that reported utility data corresponding to PTSD-related health states in children and young people that met inclusion criteria (Gospodarevskaya, 2013; Shearer 2018). No studies reporting utility data were excluded after obtaining full text.

Gospodarevskaya (2013) reported utility scores generated using HRQoL ratings of 993 adolescents and young adults aged 16 to 21 years, some of whom had a history of childhood sexual abuse prior to the age of 18, who participated in a Mental Health Survey in Australia. Diagnosis of PTSD was made using the standardized structured World Mental Health Composite International Diagnostic Interview (WMHCIDI, Version 3.0) based on the DSM-IV-TR. HRQoL was assessed with the generic Assessment of Quality of Life (AQoL) measure, which was subsequently converted to the AQoL-4D preference-based measure. The scale includes 12 items (personal care, household tasks, ability to move around the house and community, personal relationships, relationships with other people, relationships with family, vision, hearing, communication with others, sleeping habits, feelings in general, and level of pain or discomfort) rated using 4 levels. The AQoL-4D has been valued by a sample of the Australian general population using the time-trade-off (TTO) technique. The authors reported utility scores for the total study sample of adolescents and young adults (n=993); for those with history of sexual abuse prior to 18 years (n=82); for those diagnosed with PTSD due to any traumatic event (n=30); for adolescents and young adults who had PTSD associated with sexual abuse (n=14); and for those who had PTSD comorbid with depression (n=9).

Shearer and colleagues (2018) reported utility scores derived from 29 children and adolescents aged 8-17 years that met ICD-10 criteria for PTSD who participated in an 11 week RCT of early PTSD treatment (TF-CBT) versus wait list 2-6 months after a single trauma event in the UK (Meiser-Stedman 2010/2017). HRQoL was measured using the parent-completed Strengths and Difficulties Questionnaire (SDQ). SDQ scores were mapped onto the Child Health Utility index 9D (CHU-9D) using a published mapping algorithm developed in a sample of 200 young people in Australia attending child and adolescent mental health services. The CHU-9D is a generic measure of children's health state preferences consisting of nine dimensions (sad, worried, pain, annoyed, tired, homework or schoolwork, daily routine, activities and sleep) rated using five levels; preferences for this measure were elicited from the Australian population using standard gamble (SG). Baseline HRQoL data derived from all children participating in the trial were used to determine the utility of a PTSD health state. Data obtained from all children who were PTSD-free at trial follow up, irrespective of group allocation, were used to determine the utility corresponding to a PTSD-free health state.

An overview of the study characteristics, the methods used to define health states, and the health-state utility values reported by each of the two studies is provided in Table 47.

Table 47. Summary of available health-state utility data for PTSD in children and young people

| Study                    | Definition of health states  | Utility<br>measure,<br>valuation<br>method,<br>population<br>valuing          | Health states & corresponding utility  | scores  |
|--------------------------|--|---|--|---|
| Gospodarevskaya,<br>2013 | 993 adolescents and young adults aged 16-21 years, who participated in the 2007 Australian National Survey of Mental Health and Wellbeing. Diagnosis of PTSD was made using the standardized structured World Mental Health Composite International Diagnostic Interview (WMHCIDI, Version 3.0) based on the DSM-IV-TR. HRQoL was assessed with the generic Assessment of Quality of Life (AQoL) measure, which was subsequently converted to AQoL-4D. The scale includes 12 items (personal care, household tasks, ability to move around the house and community, personal relationships, relationships with other people, relationships with family, vision, hearing, communication with others, sleeping habits, feelings in general, level of pain or discomfort) rated using 4 levels. | AQoL-4D<br>TTO<br>Australian<br>general<br>population,<br>aged 16-74<br>years | Health state (Young people aged 16-21 years) General population (n=993) History of sexual abuse prior to 18 years (n=82) PTSD due to any traumatic event (n=30) PTSD associated with sexual abuse (n=14) PTSD + depression (n=9) | Mean (SD) 0.87 (0.17) 0.71 (0.25) 0.68 (0.28) 0.61 (0.31) 0.53 (0.26) |
| Shearer 2018             | 29 children and adolescents aged 8-17 years that met ICD-10 criteria for PTSD who participated in an 11 week RCT of trauma-focused CBT vs wait list 2-6 months after a single trauma event in the UK. HRQoL was measured using the parent-completed Strengths and Difficulties Questionnaire (SDQ). Data for the PTSD health state were derived from all children participating in the RCT at baseline. Data for the PTSD-free health state were obtained from all children who were PTSD-free at trial follow up  | CHU-9D<br>SG<br>UK adult<br>general<br>population                             | Health state PTSD-free following treatment (n=14) PTSD (n=29)  | Mean (95% CI)<br>0.77 (0.74-0.80)<br>0.74 (0.63-0.85)                 |

| irrespective of group allocation. SDQ scores were  | Study | Definition of health states  | Utility<br>measure,<br>valuation<br>method,<br>population<br>valuing | Health states & corresponding utility scores |
|--|-------|--|--|--|
| mapped onto the Child Health Utility index 9D (CHU- 9D) using a published mapping algorithm developed in a sample of 200 young people in Australia attending child and adolescent mental health services. CHU-9D is a generic preference-based measure for children comprising 9 dimensions (sad, worried, pain, annoyed, tired, homework or schoolwork, daily routine, activities and sleep) rated using five levels. |       | mapped onto the Child Health Utility index 9D (CHU-9D) using a published mapping algorithm developed in a sample of 200 young people in Australia attending child and adolescent mental health services. CHU-9D is a generic preference-based measure for children comprising 9 dimensions (sad, worried, pain, annoyed, tired, homework or schoolwork, daily routine, activities and sleep) rated |  |  |

CI: confidence intervals; SD: standard deviation; SG: standard gamble; TTO: time trade-off

According to NICE guidance on the selection of utility values for use in cost-utility analysis (NICE, 2013), the measurement of changes in HRQoL should be reported directly from people with the condition examined, or, if this is not possible, by their carers, and the valuation of health states should be based on public preferences elicited using a choice-based method, such as the time trade-off (TTO) or standard gamble (SG), in a representative sample of the UK population. NICE recommends the EQ-5D (Brooks, 1996; Dolan, 1997) as the preferred measure of HRQoL in adults for use in cost-utility analysis. For children, consideration should be given to alternative standardised and validated preference-based measures of health-related quality of life that have been designed specifically for use in children, as the standard version of the EQ-5D has not been designed for use in children. An alternative version for children aged 7–12 years is available, but a validated UK valuation set is not yet available.

The committee noted that the data reported by Gospodarevskaya (2013) were obtained from an overall large study sample. However, the age of the study sample (adolescents and young adults) was not directly relevant to that of the study population of the economic model, which consisted of children and young people under 18 years of age. Moreover, the utility value of the general population in the utility study was likely to be higher than the utility of a 'no PTSD health state' that was the result of remission from PTSD. It was also noted that the utility data reported by Gospodarevskaya (2013) reflected Australian population's preferences. On the other hand, data were obtained directly from participants in the survey and preferences were elicited using TTO, which meets NICE criteria for the selection of utility values.

The data reported by Shearer and colleagues (2018) were derived from a very small study sample, which, nevertheless, was directly relevant to the study population of the economic analysis (in both the utility study and the guideline economic analysis the study population was children and young people with PTSD). HRQoL ratings were obtained from parents, rather than the children themselves, and the committee considered this as a limitations as parents may easily recognise externalising / behavioural difficulties, but they may underestimate internalising symptoms (e.g. anxiety, post-traumatic stress symptoms, depression). Moreover, the committee had concerns about the ability of the SDQ instrument to capture all aspects of HRQoL and of the accuracy of mapping of these ratings onto the CHU-9D. The committee noted that the mapping algorithm was developed in a sample of Australian young people and that UK preferences were used. The committee also noted that the study population in Shearer and colleagues was at early stages of PTSD and therefore was likely to be at an early lifetime of impairment. Finally, the committee noted the narrow difference between the PTSD and the PTSD-free state utility values.

Following these considerations, the committee advised that the economic analysis utilise data from Gospodarevskaya (2013) in the base-case analysis; the values of Shearer and colleagues (2018) were utilised in sensitivity analysis.

Changes in utility between the states of 'PTSD' and 'no PTSD' were assumed to occur linearly over the time period of the change. When running the probabilistic analysis, the utility value of the 'no PTSD' health state was not allowed to become lower than that of the 'PTSD' health state. In iterations where the utility of the 'no PTSD' health state was lower than the utility of the 'PTSD' health state, the former was forced to equal the latter.

#### Intervention resource use and costs

Intervention costs were estimated by combining resource use associated with each intervention with appropriate healthcare professional unit costs.

Resource use estimates of each psychological therapy in terms of number and duration of sessions, mode of delivery and number of therapists and participants in the case of group interventions were determined by resource use data described in respective RCTs that were included in the guideline NMA that informed the economic analysis, modified by the committee to represent clinical practice in the UK. All psychological interventions were assumed to be delivered by an Agenda for Change (AfC) band 7 clinical psychologist, following expert advice from the committee on optimal delivery of psychological interventions for children and young people with PTSD. Therapist unit costs were estimated using a combination of data derived from national sources (British Association for Behavioural and Cognitive Therapies, 2016; Curtis & Burns, 2017; National College for Teaching and Leadership, NHS Health Education England, 2016) and included wages/salary, salary oncosts, capital and other overheads, qualification costs and the cost of monthly supervision. Qualification costs were annuitised using the formula reported in Netten and colleagues (1998), assuming a useful working life of 25 years, a time from obtaining the qualification until retirement of 44 years, and an equal distribution of the useful working life over the period of 44 years due to lack of specific information on this distribution. In estimating the unit cost of clinical psychologists per hour of client contact, the ratio of direct (face-to-face) to indirect time (reflecting time for preparation of therapeutic sessions and other administrative tasks) of the clinical psychologists was also taken into account.

The unit cost of a band 7 clinical psychologist was estimated to be £101 per hour of direct contact with the client. An overview of the cost elements that were taken into account in this estimation is shown in Table 48.

Table 48: Unit cost of clinical psychologist band 7 (2017 prices)

| Cost element          | Unit cost (annual) | Source   |
|-----------------------|--------------------|--|
| Wages – salary        | £38,951            |  |
| Salary on-costs       | £9,864             | Curtis & Burns, 2017; unit cost of   |
| Overheads - staff     | £11,960            | community-based scientific & professional staff, including allied health professionals   |
| Overheads - non-staff | £18,647            | (Agenda for Change band 7)   |
| Capital overheads     | £5,125             | ,  |
| Qualifications        | 12,386             | Based on a mean clinical psychologist training cost estimate of £159,420 (National College for Teaching and Leadership, NHS Health Education England, 2016), annuitised using the formula reported in Netten and colleagues (1998), assuming a useful working life of 25 years, a time from obtaining the qualification until retirement of 44 years, and an equal distribution of the useful working life over the period of 44 years due to lack of specific information on this distribution. |

| Cost element  | Unit cost (annual)                                    | Source  |  |  |
|---|---|---|--|--|
| Supervision   | £316  | Based on the unit cost of an Agenda for Change band 8a clinical psychologist (Curtis & Burns, 2017) providing 1.5 hour of supervision per month, delivered in groups of 4 participants (British Association for Behavioural and Cognitive Therapies, 2016 and expert advice); qualification costs included, as described above. |  |  |
| SUM of unit costs   | £97,249   |   |  |  |
| Working time  | 42.6 weeks /year<br>37.5 hours /week<br>(1,599 hours) | Curtis & Burns, 2017  |  |  |
| Total cost per hour   | £61   |   |  |  |
| Ratio of direct to indirect time*   | 60:40   | Curtis & Burns, 2017; assumption based on<br>the committee's expert opinion and a<br>review of respective ratios reported in the<br>literature for clinical psychologists and other<br>therapists delivering psychological<br>interventions   |  |  |
| Estimated cost per hour of direct contact   | £101  |   |  |  |
| * ratio of face-to-face time to time for preparation and other administrative tasks |   |   |  |  |

Details on the resource use and total costs of psychological interventions are provided in Table 49.

Table 49: Intervention costs of psychological therapies for children and young people with PTSD considered in the guideline economic analysis (2017 prices)

| Intervention                          | Resource use details   | Total intervention cost per person |
|---------------------------------------|--|------------------------------------|
| Supportive counselling                | 12 x 75 min individual sessions (15 hours)   | £1,520                             |
| TF-CBT (group CBT)                    | 10 x 60 min group sessions (10 hours), 1 therapist and 6 participants per group plus 1 x 60 min individual orientation meeting | £270                               |
| TF-CBT (Cohen/CPT)                    | 12 x 60 min individual/family sessions (12 hours)  | £1,216                             |
| TF-CBT (cognitive therapy)            | 10 x 90min individual sessions (15 hours)  | £1,520                             |
| TF-CBT (narrative exposure)           | 6 x 60min individual sessions (6 hours)  | £608                               |
| TF-CBT (exposure /prolonged exposure) | 14 x 60 min individual sessions (14 hours)   | £1,419                             |
| EMDR                                  | 8 x 45 min individual sessions (6 hours)   | £608                               |
| Family therapy                        | 4 x 75 min group sessions (5 hours), 1 therapist & 6 families per group plus 2 hours of individual contact                     | £287                               |

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| Intervention    | Resource use details                       | Total intervention cost per person |
|-----------------|--|------------------------------------|
| Play therapy    | 20 x 30 min individual sessions (10 hours) | £1,014                             |
| Parent training | 12 x 45 min individual sessions (9 hours)  | £912                               |
| No treatment    | No related resource use                    | £0                                 |

All interventions assumed to be delivered by a Band 7 clinical psychologist

CPT: cognitive processing therapy; EMDR: eye movement desensitisation reprocessing; TF-CBT: trauma-focused cognitive behavioural therapy

#### Costs associated with the PTSD and 'no PTSD' health states

Costs associated with the PTSD and no PTSD health states were estimated using cost data from Shearer (2018). The authors described a model-based economic analysis that utilised cost data from a 11-week RCT (Meiser-Stedman 2010/2017) that evaluated early PTSD treatment (TF-CBT) versus waitlist 2-6 months after a single trauma event in 29 children and adolescents with PTSD aged 8-17 years the UK. Cost data from the UK NHS/PSS perspective were collected for all participants at baseline and over the trial period using the Child and Adolescent Service Use Schedule (CA-SUS) and clinical records for intervention contact time. Costs included staff time (GP, nurse, paediatrician, clinical psychologist, CAMHS worker, counsellor, educational psychologist), hospital services (inpatient, outpatient, emergency department, ambulance), advice services, social services and medication. The authors estimated the total NHS/PSS cost for all children (in both arms) at baseline, and for children who were PTSD-free at trial follow up, irrespective of allocation arm (after excluding intervention cost from children in intervention arm), and extrapolated them in order to estimate annual costs incurred by children and young people with PTSD, and those without PTSD, respectively. These costs were utilised in the guideline economic analysis to express total NHS/PSS costs of the PTSD and no PTSD health states, respectively. The annual costs associated with the 'PTSD' and 'no PTSD' health states are presented in Table 50.

Table 50. Annual NHS and PSS costs incurred by children and young people with PTSD and those without PTSD (based on Shearer 2018)

| Health state | Annual NHS/PSS cost per person (2017 prices) |
|--------------|--|
| PTSD         | £2,701                                       |
| No PTSD      | £1,159                                       |

NHS and PSS costs were assumed to be the same across all arms of the economic model during the period of initial treatment (0-3 months of the economic analysis) and therefore were excluded from further consideration.

All costs were expressed in 2017 prices, uplifted, where necessary, using the Hospital and Community Health Services Pay and Prices Index (Curtis & Burns, 2017). Costs and QALYs were discounted at an annual rate of 3.5%, according to NICE guidance (NICE, 2014).

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### **Discounting**

Costs and benefits were discounted at an annual rate of 3.5% as recommended by NICE (2014).

### Handling uncertainty

Model input parameters were synthesised in a probabilistic analysis. This means that the input parameters were assigned probabilistic distributions (rather than being expressed as point estimates); this approach allowed more comprehensive consideration of the uncertainty characterising the input parameters and captured the non-linearity characterising the economic model structure. Subsequently, 10,000 iterations were performed, each drawing random values out of the distributions fitted onto the model input parameters. Results (mean costs and QALYs for each intervention) were averaged across the 10,000 iterations. This exercise provides more accurate estimates than those derived from a deterministic analysis (which utilises the mean value of each input parameter ignoring any uncertainty around the mean), by capturing the non-linearity characterising the economic model structure (Briggs 2006).

The distributions of the log-odds ratios of relative effects of all treatments versus no treatment were obtained from the respective NMAs, defined directly from values recorded in each of the 10,000 iterations used after thinning the 300,000 iterations performed in WinBUGS.

Beta distribution was assigned to the following parameters: the baseline probability of remission (probability of remission of no treatment between 0-6 months and probability of remission across all interventions from 6 months onwards); the probability of relapse; and the utility values.

The odds-ratio of dichotomous remission that was applied to all active interventions versus waitlist was assigned a log-normal distribution.

Uncertainty in intervention costs was taken into account by assigning probability distributions to the number of individually delivered psychological therapy sessions, based on intervention completion data and data on mean number of sessions reported in the RCTs that informed the economic analysis. The number of therapist sessions per person attending group psychological interventions was not assigned a probability distribution because the number of group sessions remains the same, whether a participant attends the full course of treatment or a lower number of sessions. The unit cost of clinical psychologists was assigned a normal distribution.

NHS/PSS costs associated with the 'PTSD' and 'no PTSD' health states were assigned a gamma distribution.

Table 51 reports the mean values of all input parameters utilised in the economic model and provides details on the types of distributions assigned to each input parameter and the methods employed to define their range.

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

Table 51: Input parameters (deterministic values and probability distributions) that informed the economic model of interventions for the treatment of PTSD in children and young people

| Input parameter   | Mean<br>deterministic<br>value | Probability distribution | Source of data - comments                                    |  |
|---|--------------------------------|--------------------------|--|--|
| Odds ratios of remission versus no trea   | atment/waitlist a              | at treatment endpoint    |  |  |
| Derived from NMA of continuous data   |                                | 95% Crl                  |  |  |
| Supportive counselling  | 2.97                           | 0.84 to 10.64            |  |  |
| TF-CBT (group CBT)  | 5.21                           | 1.87 to 14.60            |  |  |
| TF-CBT (Cohen/CPT)  | 8.43                           | 2.74 to 26.05            |  |  |
| TF-CBT (CT)   | 204.50                         | 34.36 to 1271.56         |  |  |
| TF-CBT (narrative exposure)   | 15.14                          | 3.99 to 59.20            |  |  |
| TF-CBT (exposure /prolonged exposure)   | 11.42                          | 2.65 to 50.55            |  |  |
| EMDR  | 6.09                           | 1.52 to 24.80            |  |  |
| Family therapy  | 1.96                           | 0.22 to 19.03            | Guideline NMA; distribution based on 10,000 iterations       |  |
| Play therapy  | 11.52                          | 1.51 to 90.65            |  |  |
| Parent training   | 5.83                           | 0.49 to 66.95            |  |  |
| Derived from NMA of remission data  |                                |                          |  |  |
| Supportive counselling  | 1.16                           | 0.37 to 3.56             |  |  |
| TF-CBT (CT)   | 14.29                          | 3.65 to 66.09            |  |  |
| TF-CBT (Cohen/CPT)  | 2.44                           | 1.16 to 5.08             |  |  |
| TF-CBT (narrative exposure)   | 16.71                          | 2.38 to 176.80           |  |  |
| TF-CBT (exposure /prolonged exposure)   | 5.07                           | 1.24 to 20.66            |  |  |
| Odds ratios of remission versus no treatment/waitlist at 3-month follow-up (sensitivity analysis) |                                |                          |  |  |
| Derived from NMA of continuous data   |                                | 95% Crl                  | Guideline NMA; distribution based on 10,000 iterations       |  |
| Supportive counselling  | 3.83                           | 0.89 to 12.99            | Estimated 3-6 month probability of remission for CT borrowed |  |
| TF-CBT (group CBT)  | 15.51                          | 2.90 to 91.56            | from Cohen/CPT; estimated 3-6 months probability of          |  |
| TF-CBT (Cohen/CPT)  | 23.82                          | 2.19 to 285.43           | remission for family therapy and play therapy assumed to     |  |
| TF-CBT (CT)   | No data                        | No data                  | equal that of no treatment                                   |  |

## DRAFT FOR CONSULTATION

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Input parameter                             | Mean<br>deterministic<br>value | Probability distribution          | Source of data - comments  |
|---|--------------------------------|-----------------------------------|--|
| TF-CBT (narrative exposure)                 | 5.54                           | 1.09 to 28.05                     |  |
| TF-CBT (exposure/prolonged exposure)        | 5.31                           | 0.48 to 57.80                     |  |
| EMDR  | 2.94                           | 0.18 to 47.13                     |  |
| Parent training                             | 6.51                           | 0.23 to 197.35                    |  |
| Family therapy                              | No data                        | No data                           |  |
| Play therapy                                | No data                        | No data                           |  |
| Derived from pairwise meta-analysis of      |                                | Log-normal distribution:          | Guideline pairwise meta-analysis   |
| <u>dichotomous data – all interventions</u> | 25.86                          | 95% CI 7.28 to 91.84              |  |
| Probability of remission – no treatment     | t                              |                                   |  |
| 0-3 months from PTSD onset                  | 0.174                          | Beta: α=87.00; β=413.00           | Rosellini 2017; data averaged between children aged 0-12   |
| 0-12 months from PTSD onset                 | 0.370                          | Beta: α=185.19; β=314.81          | years and young people aged 13-24 years; 3-month   |
| 0-24 months from PTSD onset                 | 0.445                          | Beta: α=222.26; β=277.74          | probabilities estimated using the cumulative remission data  |
| 0-36 months from PTSD onset                 | 0.500                          | Beta: α=250.00; β=250.00          | after excluding the first 3 months from PTSD onset as the model study population received treatment after 3 months from PTSD onset |
| Risk of relapse – all model arms            |                                |                                   |  |
| 3-month risk                                | 0.026                          | Beta: α=2.60; β=97.40             | Assumption   |
| Utility values                              |                                | Beta distribution                 |  |
| Base-case analysis                          |                                |                                   |  |
| PTSD – 3-month                              | 0.170                          | $\alpha$ =9.01; $\beta$ =43.98    | Gospodarevskaya, 2013; distribution estimated based on   |
| No PTSD – 3-month                           | 0.218                          | α=1271.69; β=4575.15              | method of moments  |
| Sensitivity analysis                        |                                |                                   |  |
| PTSD – 3-month                              | 0.185                          | α=808; β=3,567                    | Shearer 2018   |
| No PTSD – 3-month                           | 0.193                          | α=2,618; β=10,940                 |  |
| Intervention costs - resource use           |                                |                                   | Probabilities assigned to numbers of sessions  |
| Number of sessions                          |                                |                                   |  |
| Supportive counselling                      | 12                             | 0.60: 10-12, 0.22: 6-9, 0.18: 3-5 | Number of visits and probabilities based on resource use and   |
| TF-CBT (group CBT)                          | 10                             | No distribution                   | completion rate data reported in the RCTs included in the  |

## DRAFT FOR CONSULTATION

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Input parameter                           | Mean<br>deterministic<br>value | Probability distribution                     | Source of data - comments   |
|---|--------------------------------|--|---|
| TF-CBT (Cohen/CPT)                        | 12                             | 0.60: 10-12, 0.22: 6-9, 0.18: 3-5            | NMAs that informed the economic analysis  |
| TF-CBT (CT)                               | 10                             | 0.70: 8-10, 0.16: 6-7, 0.14: 3-5             |   |
| TF-CBT (narrative exposure)               | 6                              | 0.80: 5-6, 0.10: 4, 0.10: 3                  |   |
| TF-CBT (exposure/prolonged exposure)      | 14                             | 0.70: 11-14, 0.16: 7-10, 0.14: 3-6           |   |
| EMDR                                      | 8                              | 0.60: 7-8, 0.22: 4-6, 0.18: 2-3              |   |
| Family therapy                            | 4                              | No distribution                              |   |
| Play therapy                              | 20                             | 0.60: 14-20, 0.22: 10-13, 0.18: 7-9          |   |
| Parent training                           | 12                             | 0.60: 10-12, 0.22: 6-9, 0.18: 3-5            |   |
| Unit cost of clinical psychologist Band 7 | £101                           | Normal distribution<br>SE = 0.05 of the mean | Estimated using data from the British Association for Behavioural and Cognitive Therapies, 2016; Curtis & Burns, 2017; National College for Teaching and Leadership, NHS Health Education England, 2016; distribution based on assumption |
| 3-month NHS/PSS health state cost         |                                | Gamma distribution                           |   |
| PTSD                                      | £549                           | α=19.53; β=28.12                             | Shearer (2018), expressed in 2017 prices using the HCHS   |
| No PTSD                                   | £236                           | α=10.37; β=22.74                             | inflation index (Curtis & Burns, 2017).   |
| Annual discount rate                      | 0.035                          | No distribution                              | Applied to both costs and outcomes (NICE, 2014)   |

A number of different scenarios were explored by using the 2 sets of available utility data and 2 alternative assumptions on the efficacy of interventions at the 3-month follow-up. Consequently, 4 separate probabilistic analyses were undertaken:

- Scenario A: Use of utility data derived from Gospodarevskaya (2013); the probability of remission of all active interventions at 3-6 months was conservatively assumed to be equal to that of no treatment. This scenario formed the base-case economic analysis.
- Scenario B: Use of utility data derived from Gospodarevskaya (2013); the relative effect
  of active interventions versus no treatment at 3-6 months was derived from the NMA of
  changes in PTSD symptom scores between baseline and 1-4 month follow-up.
- Scenario C: Use of utility data derived from Shearer (2018); the probability of remission of all active interventions at 3-6 months was assumed to be equal to that of no treatment.
- Scenario D: Use of utility data derived from Shearer (2018); the relative effect of active
  interventions versus no treatment at 3-6 months was derived from the NMA of changes in
  PTSD symptom scores between baseline and 1-4 month follow-up.

For all these scenarios, secondary analysis was run in parallel, which utilised the output of the NMA of dichotomous remission data at treatment endpoint. For scenarios (B) and (D) the odds ratio of group CBT versus waitlist at 1-4 month follow-up was used to estimate the relative effect of all interventions versus no treatment at 3-6 months.

One-way deterministic sensitivity analysis was also employed to explore the impact of a change in the annual risk of relapse, which was varied between zero and 0.20.

#### Presentation of the results

Results of the economic analysis are presented as follows:

In each analysis, mean total costs and QALYs are presented for each intervention, averaged across 10,000 iterations of the model. An incremental analysis is provided for each cohort, in table format, where all options have been listed from the most to the least effective (in terms of QALYs gained). Options that are dominated by absolute dominance (that is, they are less effective and more costly than one or more other options) or by extended dominance (that is, they are less effective and more costly than a linear combination of two alternative options) are excluded from further analysis. Subsequently, incremental cost-effectiveness ratios (ICERs) are calculated for all pairs of consecutive options remaining in analysis.

ICERs are calculated by the following formula:

ICER = 
$$\Delta C / \Delta E$$

where  $\Delta C$  is the difference in total costs between two interventions and  $\Delta E$  the difference in their effectiveness (QALYs). ICERs express the extra cost per extra unit of benefit (QALY) associated with one treatment option relative to its comparator. The treatment option with the highest ICER below the NICE lower cost effectiveness threshold of £20,000/QALY (NICE 2008) is the most cost-effective option.

In addition to ICERs, the mean net monetary benefit (NMB) of each intervention is presented. This is defined by the following formula:

NMB = 
$$\mathbf{E} \cdot \lambda - \mathbf{C}$$

where E and C are the effectiveness (number of QALYs) and costs associated with the treatment option, respectively, and  $\lambda$  is the level of the willingness-to-pay (WTP) per unit of effectiveness, set at the NICE lower cost effectiveness threshold of £20,000/QALY (NICE, 2008). The intervention with the highest NMB is the most cost-effective option (Fenwick 2001).

Incremental mean costs and effects (QALYs) of each intervention versus no treatment are also presented in the form of cost effectiveness planes.

The probability of each intervention being the most cost-effective option at the NICE lower cost effectiveness threshold of £20,000/QALY is provided, calculated as the proportion of iterations (out of the 10,000 iterations run) in which the intervention has had the highest NMB among all interventions considered in the analysis.

The mean ranking in terms of cost effectiveness is also reported for each intervention (out of the 10,000 iterations run), where a rank of 1 is best.

The probabilities of each intervention being cost-effective at various cost effectiveness thresholds are illustrated in cost-effectiveness acceptability curves (CEACs). Finally, the cost-effectiveness acceptability frontiers (CEAFs) are also plotted; these show the treatment option with the highest mean NMB over different cost effectiveness thresholds, and the probability that the option with the highest NMB is the most cost-effective among those assessed (Fenwick 2001).

#### Validation of the economic model

The economic model (including the conceptual model and the identification and selection of input parameters) was developed by the health economist in collaboration with a health economics sub-group formed by members of the committee. As part of the model validation, all inputs and model formulae were systematically checked; the model was tested for logical consistency by setting input parameters to null and extreme values and examining whether results changed in the expected direction. The base-case results and results of sensitivity analyses were discussed with the committee to confirm their plausibility.

## **Economic modelling results**

## Scenario A, base-case analysis: utility data from Gospodarevskaya (2013); no beneficial effect beyond treatment endpoint

The results of the base-case economic analysis are provided in Table 52. This table provides mean QALYs and mean total costs for each intervention assessed in the economic analysis, as well as the results of incremental analysis, the mean NMB of each intervention, and its mean ranking by cost effectiveness (where a rank of 1 is best). Interventions have been ordered from the most to the least effective in terms of number of QALYs gained. According to the results, cognitive therapy was the most clinically and cost-effective intervention, followed by narrative exposure and play therapy. Prolonged exposure and Cohen/CPT were in the top 5 most clinically and cost-effective treatment options. All

interventions were more effective and cost-effective than no treatment. In the secondary analysis that utilised dichotomous remission data, all forms of TF-CBT were more effective and cost-effective than no treatment; however, supportive counselling was less cost-effective than no treatment.

Table 52: Scenario A, base-case results of economic modelling: interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); no beneficial effect beyond treatment endpoint]

| Intervention   | Mean per person |              |              | ICER      | NMB £/ | Prob              | Mean |
|--|-----------------|--------------|--------------|-----------|--------|-------------------|------|
|  | QALY            | Inter cost £ | Total cost £ | (£/QALY)  | person | best <sup>1</sup> | rank |
| Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores |                 |              |              |           |        |                   |      |
| TF-CBT cognitive therapy   | 2.467           | 1,202        | 4,347        | Dominant  | 44,993 | 0.78              | 1.57 |
| TF-CBT narrative exposure  | 2.322           | 517          | 4,484        | Dominated | 41,966 | 0.08              | 3.35 |
| TF-CBT prolonged exposure  | 2.297           | 1,089        | 5,200        | Dominated | 40,742 | 0.01              | 5.35 |
| Play therapy   | 2.297           | 719          | 4,827        | Dominated | 41,109 | 0.05              | 4.68 |
| TF-CBT Cohen/CPT   | 2.268           | 915          | 5,188        | Dominated | 40,178 | 0.00              | 5.91 |
| Parent training  | 2.244           | 684          | 5,099        | Dominated | 39,788 | 0.03              | 6.50 |
| EMDR   | 2.241           | 460          | 4,897        | Dominated | 39,920 | 0.01              | 5.88 |
| TF-CBT group CBT   | 2.224           | 270          | 4,798        | Dominated | 39,687 | 0.01              | 5.83 |
| Supportive counselling   | 2.183           | 1,141        | 5,902        | Dominated | 37,753 | 0.00              | 9.57 |
| Family therapy   | 2.168           | 287          | 5,133        | Dominated | 38,222 | 0.01              | 8.20 |
| No treatment   | 2.121           | 0            | 5,113        | Dominated | 37,304 | 0.01              | 9.16 |
| Analysis utilising efficacy data from NMAs of dichotomous remission          |                 |              |              |           |        |                   |      |
| TF-CBT narrative exposure  | 2.326           | 517          | 4,464        | Dominant  | 42,066 | 0.62              | 1.53 |
| TF-CBT cognitive therapy   | 2.317           | 1,202        | 5,203        | Dominated | 41,128 | 0.30              | 2.10 |
| TF-CBT prolonged exposure  | 2.225           | 1,089        | 5,612        | Dominated | 38,888 | 0.05              | 3.27 |
| TF-CBT Cohen/CPT   | 2.165           | 915          | 5,777        | Dominated | 37,525 | 0.00              | 4.29 |
| Supportive counselling   | 2.131           | 1,141        | 6,198        | Dominated | 36,418 | 0.00              | 5.73 |
| No treatment   | 2.121           | 0            | 5,113        | Dominated | 37,304 | 0.03              | 4.09 |

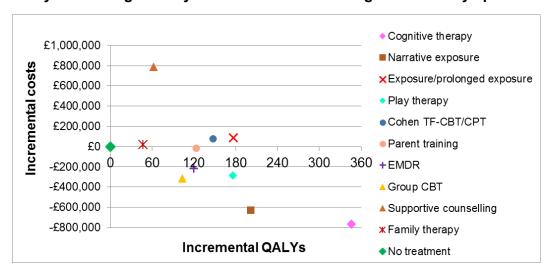
<sup>1</sup> at the NICE lower cost-effectiveness threshold of £20,000/QALY

CPT: cognitive processing therapy; EMDR: eye movement desensitisation reprocessing; ICER: incremental cost effectiveness ratio; Inter: intervention; NMB: net monetary benefit; Prob: probability; TF-CBT: trauma-focused cognitive behavioural therapy

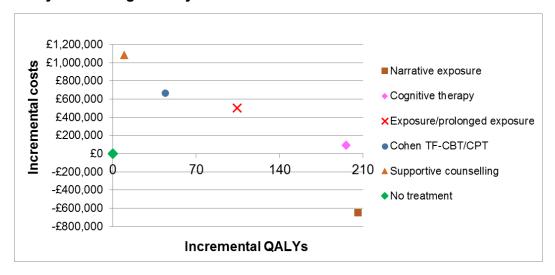
Figure 176 provides the cost effectiveness plane of the analysis. Each intervention is placed on the plane according to its incremental costs and QALYs compared with no treatment, which is placed at the origin.

Figure 176. Scenario A, base-case analysis: Cost-effectiveness plane of interventions for the treatment of PTSD in children and young people plotted against no treatment – incremental costs and QALYs per 1,000 children and young people [utility data from Gospodarevskaya (2013); no beneficial effect beyond treatment endpoint]

#### Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores



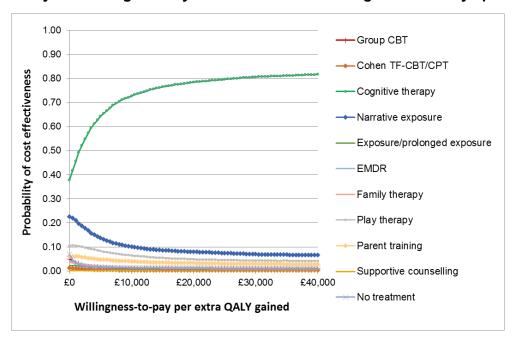
### Analysis utilising efficacy data from NMAs of dichotomous remission



The CEAC and CEAF of the analysis are shown in Figure 177 and Figure 178, respectively. It can be seen that cognitive therapy (analysis utilising NMA of continuous data) and narrative exposure (analysis utilising NMA of dichotomous data) are the most cost-effective options at any cost effectiveness threshold between zero and £40,000/QALY, with a probability that exceeds 0.60 at the NICE lower cost effectiveness threshold of £20,000/QALY.

Figure 177. Scenario A, base-case analysis: Cost-effectiveness acceptability curves of interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); no beneficial effect beyond treatment endpoint]

#### Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores



### Analysis utilising efficacy data from NMAs of dichotomous remission

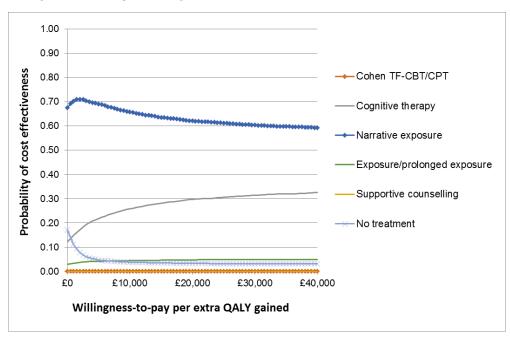
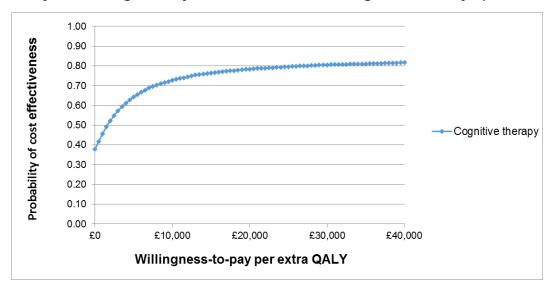
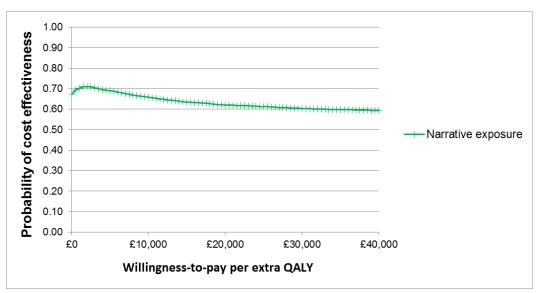


Figure 178 Scenario A, base-case analysis: Cost-effectiveness acceptability frontier of interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); no beneficial effect beyond treatment endpoint]

## Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores



## Analysis utilising efficacy data from NMAs of dichotomous remission



Results were robust to changes in the annual risk of relapse between zero and 0.20 and conclusions on cost effectiveness were not affected.

# Scenario B: utility data from Gospodarevskaya (2013); beneficial effect up to 3-month follow-up

The results of this scenario are provided in Table 53. Cognitive therapy was the most clinically and cost-effective intervention, followed by Cohen/CPT and group CBT. Narrative exposure and parent training were also among the 5 most clinically and cost-effective treatment options. All interventions were more effective and cost-effective than no treatment. In the secondary analysis that utilised dichotomous remission data, narrative exposure was the most clinically and cost-effective intervention and all interventions were more effective and cost-effective than no treatment.

Table 53: Scenario B, results of economic modelling: interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); beneficial effect up to 3-month follow-up]

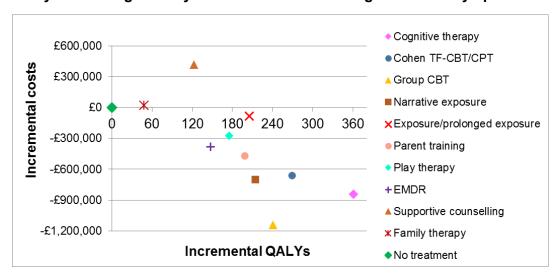
|  | Mea       | n per per       | son          | ICER          | NMB £/ | Duah                      | Magn         |  |  |
|--|-----------|-----------------|--------------|---------------|--------|---------------------------|--------------|--|--|
| Intervention   | QALY      | Inter<br>cost £ | Total cost £ | (£/QALY)      | person | Prob<br>best <sup>1</sup> | Mean<br>rank |  |  |
| Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores |           |                 |              |               |        |                           |              |  |  |
| TF-CBT cognitive therapy   | 2.482     | 1,204           | 4,271        | 2,497         | 45,373 | 0.67                      | 1.88         |  |  |
| TF-CBT Cohen/CPT   | 2.390     | 911             | 4,453        | Ext domin     | 43,348 | 0.05                      | 3.90         |  |  |
| TF-CBT group CBT   | 2.362     | 270             | 3,971        |               | 43,269 | 0.15                      | 3.35         |  |  |
| TF-CBT narrative exposure  | 2.335     | 517             | 4,414        | Dominated     | 42,296 | 0.02                      | 4.71         |  |  |
| TF-CBT prolonged exposure  | 2.326     | 1,089           | 5,033        | Dominated     | 41,495 | 0.01                      | 6.26         |  |  |
| Parent training  | 2.320     | 685             | 4,645        | Dominated     | 41,751 | 0.05                      | 5.47         |  |  |
| Play therapy   | 2.297     | 719             | 4,840        | Dominated     | 41,094 | 0.02                      | 6.31         |  |  |
| EMDR   | 2.268     | 461             | 4,731        | Dominated     | 40,636 | 0.02                      | 6.65         |  |  |
| Supportive counselling   | 2.244     | 1,135           | 5,534        | Dominated     | 39,341 | 0.00                      | 8.61         |  |  |
| Family therapy   | 2.169     | 287             | 5,135        | Dominated     | 38,245 | 0.00                      | 9.12         |  |  |
| No treatment   | 2.121     | 0               | 5,114        | Dominated     | 37,312 | 0.01                      | 9.76         |  |  |
| Analysis utilising efficacy da   | ta from N | IMAs of d       | ichotomo     | ous remission | 1      |                           |              |  |  |
| TF-CBT narrative exposure  | 2.415     | 517             | 3,934        | Dominant      | 44,370 | 0.81                      | 1.23         |  |  |
| TF-CBT cognitive therapy   | 2.411     | 1,204           | 4,647        | Dominated     | 43,570 | 0.10                      | 2.97         |  |  |
| TF-CBT prolonged exposure  | 2.397     | 1,089           | 4,581        | Dominated     | 43,349 | 0.04                      | 3.39         |  |  |
| TF-CBT Cohen/CPT   | 2.389     | 911             | 4,428        | Dominated     | 43,343 | 0.03                      | 3.30         |  |  |
| Supportive counselling   | 2.384     | 1,135           | 4,665        | Dominated     | 43,016 | 0.01                      | 4.35         |  |  |
| No treatment   | 2.121     | 0               | 5,114        | Dominated     | 37,312 | 0.01                      | 5.77         |  |  |

<sup>1</sup> at the NICE lower cost-effectiveness threshold of £20,000/QALY

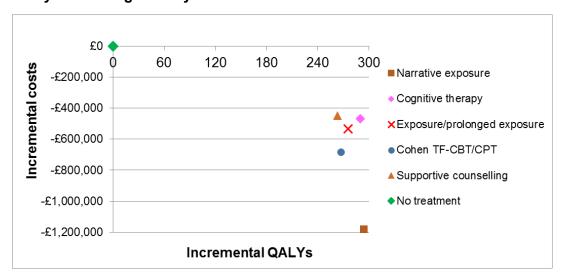
CPT: cognitive processing therapy; EMDR: eye movement desensitisation reprocessing; Ext domin: extendedly dominated; ICER: incremental cost effectiveness ratio; Inter: intervention; NMB: net monetary benefit; Prob: probability; TF-CBT: trauma-focused cognitive behavioural therapy

Figure 179 provides the cost effectiveness plane of the analysis. Each intervention is placed on the plane according to its incremental costs and QALYs compared with no treatment.

Figure 179. Scenario B: Cost-effectiveness plane of interventions for the treatment of PTSD in children and young people plotted against no treatment – incremental costs and QALYs per 1,000 children and young people [utility data from Gospodarevskaya (2013); beneficial effect up to 3-month follow-up]

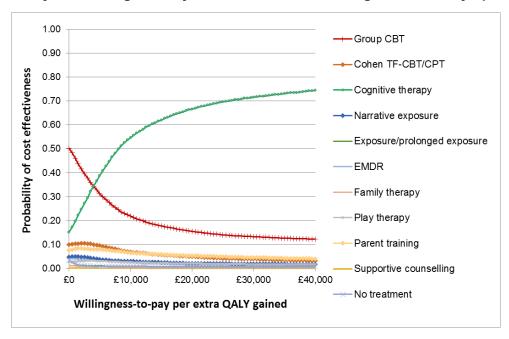


#### Analysis utilising efficacy data from NMAs of dichotomous remission



The CEAC and CEAF of the analysis are shown in Figure 180 and Figure 181, respectively. Cognitive therapy (analysis utilising NMA of continuous data) and narrative exposure (analysis utilising NMA of dichotomous data) are the most cost-effective options at the NICE lower cost effectiveness threshold of £20,000/QALY, each with a high probability of being cost-effective that exceeds 0.65.

Figure 180. Scenario B: Cost-effectiveness acceptability curves of interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); beneficial effect up to 3-month follow-up]



### Analysis utilising efficacy data from NMAs of dichotomous remission

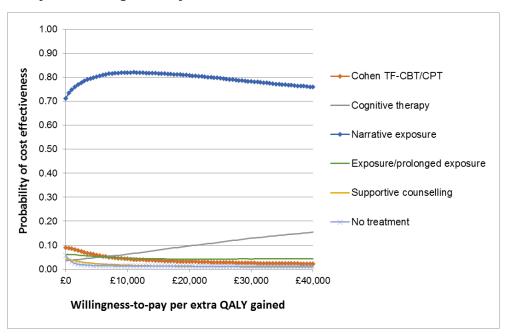
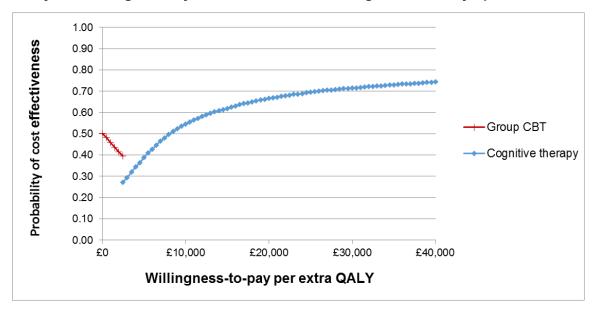
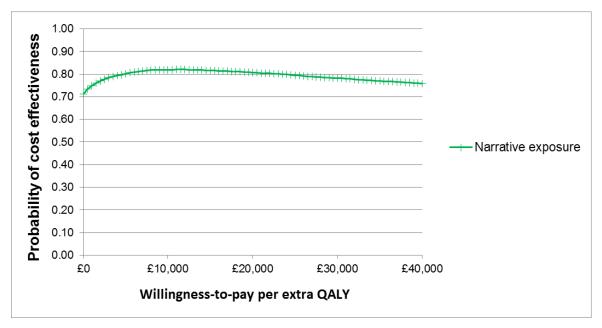


Figure 181 Scenario B: Cost-effectiveness acceptability frontier of interventions for the treatment of PTSD in children and young people [utility data from Gospodarevskaya (2013); beneficial effect up to 3-month follow-up]



## Analysis utilising efficacy data from NMAs of dichotomous remission



Results were robust to changes in the annual risk of relapse between zero and 0.20 and conclusions on cost effectiveness were not affected.

# Scenario C: utility data from Shearer (2018); no beneficial effect beyond treatment endpoint

The results of this scenario are provided in Table 54. Cognitive therapy remained the most cost-effective intervention, followed by narrative exposure and play therapy. Group CBT and EMDR were the 4<sup>th</sup> and 5<sup>th</sup> most cost-effective treatment options. All interventions were more clinically and cost-effective than no treatment, with the exception of supportive counselling, which was more clinically effective but less cost-effective. In the secondary analysis that utilised dichotomous remission data, narrative exposure followed by cognitive therapy were more cost-effective than no treatment; however, all other interventions were less cost-effective than no treatment.

Table 54: Scenario C, results of economic modelling: interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); no beneficial effect beyond treatment endpoint]

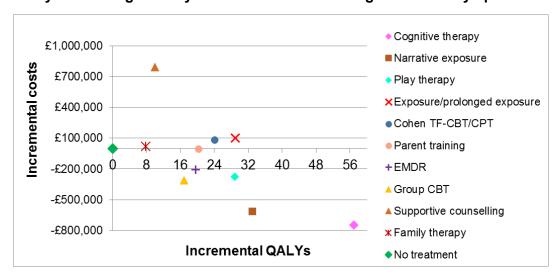
|  | Mea                       | ın per per | son      | ICED          | NMB £/                    | Duck         | Maan  |  |  |
|--|---------------------------|------------|----------|---------------|---------------------------|--------------|-------|--|--|
| Intervention   | QALY Inter Total (£/QALY) |            |          | person        | Prob<br>best <sup>1</sup> | Mean<br>rank |       |  |  |
| Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores |                           |            |          |               |                           |              |       |  |  |
| TF-CBT cognitive therapy   | 2.224                     | 1,203      | 4,373    | Dominant      | 40,108                    | 0.59         | 2.05  |  |  |
| TF-CBT narrative exposure  | 2.200                     | 517        | 4,502    | Dominated     | 39,501                    | 0.16         | 3.11  |  |  |
| TF-CBT prolonged exposure  | 2.196                     | 1,089      | 5,221    | Dominated     | 38,700                    | 0.02         | 6.47  |  |  |
| Play therapy   | 2.196                     | 715        | 4,843    | Dominated     | 39,075                    | 0.09         | 4.85  |  |  |
| TF-CBT Cohen/CPT   | 2.191                     | 911        | 5,202    | Dominated     | 38,622                    | 0.01         | 6.66  |  |  |
| Parent training  | 2.187                     | 682        | 5,112    | Dominated     | 38,635                    | 0.05         | 6.61  |  |  |
| EMDR   | 2.187                     | 459        | 4,908    | Dominated     | 38,824                    | 0.02         | 5.59  |  |  |
| TF-CBT group CBT   | 2.184                     | 270        | 4,807    | Dominated     | 38,872                    | 0.02         | 5.05  |  |  |
| Supportive counselling   | 2.177                     | 1,137      | 5,911    | Dominated     | 37,631                    | 0.00         | 10.21 |  |  |
| Family therapy   | 2.175                     | 287        | 5,139    | Dominated     | 38,357                    | 0.02         | 7.59  |  |  |
| No treatment   | 2.167                     | 0          | 5,118    | Dominated     | 38,224                    | 0.02         | 7.82  |  |  |
| Analysis utilising efficacy da   | ta from N                 | IMAs of d  | ichotomo | ous remission |                           |              |       |  |  |
| TF-CBT narrative exposure  | 2.201                     | 517        | 4,485    | Dominant      | 39,529                    | 0.71         | 1.41  |  |  |
| TF-CBT cognitive therapy   | 2.199                     | 1,203      | 5,219    | Dominated     | 38,766                    | 0.19         | 2.52  |  |  |
| TF-CBT prolonged exposure  | 2.184                     | 1,089      | 5,625    | Dominated     | 38,056                    | 0.04         | 3.76  |  |  |
| TF-CBT Cohen/CPT   | 2.174                     | 911        | 5,783    | Dominated     | 37,703                    | 0.00         | 4.58  |  |  |
| Supportive counselling   | 2.169                     | 1,137      | 6,200    | Dominated     | 37,174                    | 0.00         | 5.68  |  |  |
| No treatment   | 2.167                     | 0          | 5,118    | Dominated     | 38,224                    | 0.05         | 3.06  |  |  |

<sup>1</sup> at the NICE lower cost-effectiveness threshold of £20,000/QALY

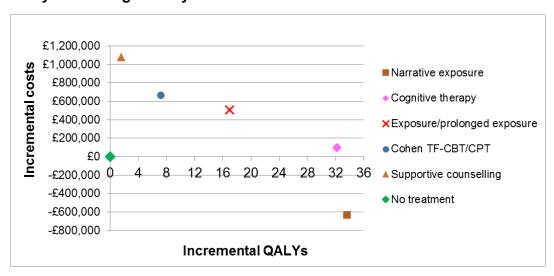
CPT: cognitive processing therapy; EMDR: eye movement desensitisation reprocessing; ICER: incremental cost effectiveness ratio; Inter: intervention; NMB: net monetary benefit; Prob: probability; TF-CBT: trauma-focused cognitive behavioural therapy

Figure 182 provides the cost effectiveness plane of the analysis. Each intervention is placed on the plane according to its incremental costs and QALYs compared with no treatment.

Figure 182. Scenario C: Cost-effectiveness plane of interventions for the treatment of PTSD in children and young people plotted against no treatment – incremental costs and QALYs per 1,000 children and young people [utility data from Shearer (2018); no beneficial effect beyond treatment endpoint]

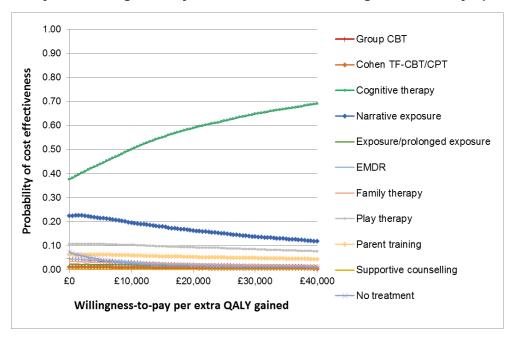


### Analysis utilising efficacy data from NMAs of dichotomous remission



The CEAC and CEAF of the analysis are shown in Figure 183 and Figure 184, respectively. Cognitive therapy (analysis utilising NMA of continuous data) and narrative exposure (analysis utilising NMA of dichotomous data) are the most cost-effective options at any cost effectiveness threshold between zero and £40,000/QALY, with a probability of being cost-effective above 0.55 at the NICE lower cost effectiveness threshold of £20,000/QALY.

Figure 183. Scenario C: Cost-effectiveness acceptability curves of interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); no beneficial effect beyond treatment endpoint]



## Analysis utilising efficacy data from NMAs of dichotomous remission

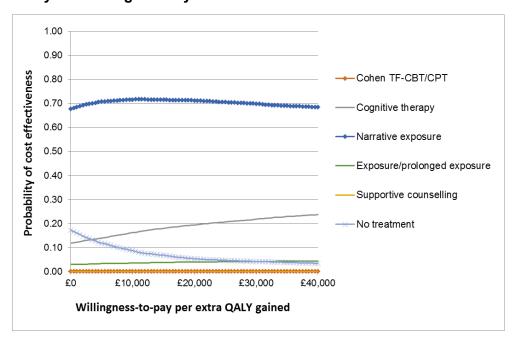
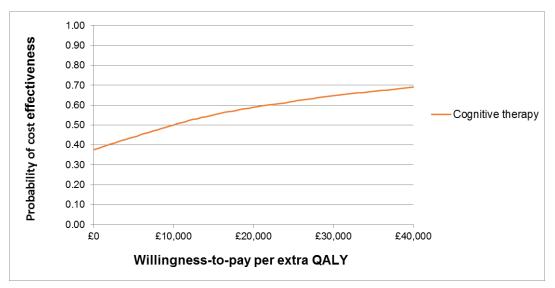
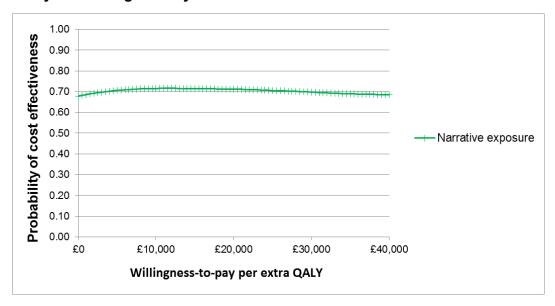


Figure 184 Scenario C: Cost-effectiveness acceptability frontier of interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); no beneficial effect beyond treatment endpoint]



## Analysis utilising efficacy data from NMAs of dichotomous remission



Results were sensitive to an increase in the annual risk of relapse from 0.10 to 0.20, as only cognitive therapy, narrative exposure, group CBT, play therapy and EMDR remained more cost-effective than no treatment. Results were not affected when a zero risk of relapse was assumed.

# Scenario D: utility data derived from Shearer (2018); beneficial effect up to 3-month follow-up

The results of this scenario are provided in Table 55. Cognitive therapy is the most cost-effective intervention in this scenario as well, followed by group CBT and Cohen/CPT. Narrative exposure and parent training were the 4<sup>th</sup> and 5<sup>th</sup> most cost-effective options, respectively. All interventions were more clinically and cost-effective than no treatment, with the exception of supportive counselling, which was more clinically effective but less cost-effective. In the secondary analysis that utilised dichotomous remission data, narrative exposure followed by Cohen/CPT and prolonged exposure were the 3 most cost-effective treatment options. All interventions were more cost-effective compared with no treatment.

Table 55: Scenario D, results of economic modelling: interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); beneficial effect up to 3-month follow-up]

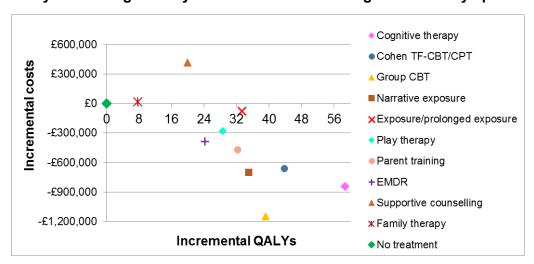
|                                | Mean per person  |              | ICER         | NMB £/        | Drob   | Moon                      |              |  |  |  |
|--------------------------------|--|--------------|--------------|---------------|--------|---------------------------|--------------|--|--|--|
| Intervention                   | QALY   | Inter cost £ | Total cost £ | (£/QALY)      | person | Prob<br>best <sup>1</sup> | Mean<br>rank |  |  |  |
| Analysis utilising efficacy da | Analysis utilising efficacy data from NMAs of changes in PTSD symptom scores |              |              |               |        |                           |              |  |  |  |
| TF-CBT cognitive therapy       | 2.227  | 1,203        | 4,271        | 15,627        | 40,276 | 0.31                      | 2.79         |  |  |  |
| TF-CBT Cohen/CPT               | 2.212  | 910          | 4,452        | Ext domin     | 39,798 | 0.10                      | 4.26         |  |  |  |
| TF-CBT group CBT               | 2.208  | 270          | 3,966        |               | 40,190 | 0.37                      | 2.54         |  |  |  |
| TF-CBT narrative exposure      | 2.204  | 518          | 4,412        | Dominated     | 39,661 | 0.05                      | 4.40         |  |  |  |
| TF-CBT prolonged exposure      | 2.202  | 1,087        | 5,035        | Dominated     | 39,004 | 0.01                      | 7.02         |  |  |  |
| Parent training                | 2.201  | 681          | 4,642        | Dominated     | 39,376 | 0.08                      | 5.45         |  |  |  |
| Play therapy                   | 2.197  | 718          | 4,833        | Dominated     | 39,113 | 0.03                      | 6.40         |  |  |  |
| EMDR                           | 2.193  | 462          | 4,727        | Dominated     | 39,130 | 0.03                      | 6.25         |  |  |  |
| Supportive counselling         | 2.189  | 1,136        | 5,529        | Dominated     | 38,244 | 0.00                      | 9.32         |  |  |  |
| Family therapy                 | 2.176  | 287          | 5,132        | Dominated     | 38,395 | 0.01                      | 8.68         |  |  |  |
| No treatment                   | 2.169  | 0            | 5,113        | Dominated     | 38,261 | 0.01                      | 8.88         |  |  |  |
| Analysis utilising efficacy da | ta from N  | IMAs of d    | ichotomo     | ous remission | 1      |                           |              |  |  |  |
| TF-CBT narrative exposure      | 2.217  | 518          | 3,929        | Dominant      | 40,404 | 0.79                      | 1.23         |  |  |  |
| TF-CBT cognitive therapy       | 2.216  | 1,203        | 4,640        | Dominated     | 39,678 | 0.04                      | 3.62         |  |  |  |
| TF-CBT prolonged exposure      | 2.214  | 1,087        | 4,572        | Dominated     | 39,699 | 0.06                      | 3.49         |  |  |  |
| TF-CBT Cohen/CPT               | 2.212  | 910          | 4,420        | Dominated     | 39,826 | 0.07                      | 2.93         |  |  |  |
| Supportive counselling         | 2.212  | 1,136        | 4,660        | Dominated     | 39,572 | 0.03                      | 4.08         |  |  |  |
| No treatment                   | 2.169  | 0            | 5,113        | Dominated     | 38,261 | 0.01                      | 5.63         |  |  |  |

<sup>1</sup> at the NICE lower cost-effectiveness threshold of £20,000/QALY

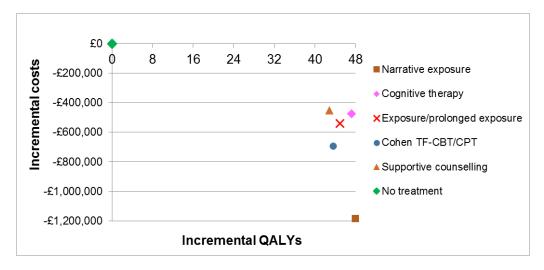
CPT: cognitive processing therapy; EMDR: eye movement desensitisation reprocessing; Ext domin: extendedly dominated; ICER: incremental cost effectiveness ratio; Inter: intervention; NMB: net monetary benefit; Prob: probability; TF-CBT: trauma-focused cognitive behavioural therapy

Figure 185 provides the cost effectiveness plane of the analysis. Each intervention is placed on the plane according to its incremental costs and QALYs compared with no treatment.

Figure 185. Scenario D: Cost-effectiveness plane of interventions for the treatment of PTSD in children and young people plotted against no treatment – incremental costs and QALYs per 1,000 children and young people [utility data from Shearer (2018); beneficial effect up to 3-month follow-up]

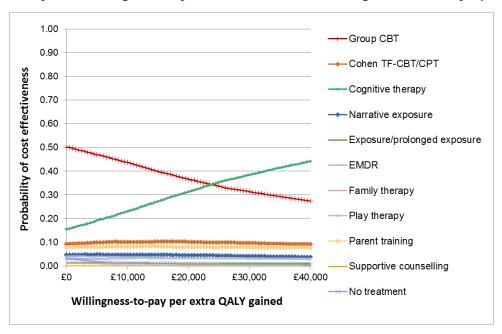


## Analysis utilising efficacy data from NMAs of dichotomous remission



The CEAC and CEAF of the analysis are shown in Figure 186 and Figure 187, respectively. Cognitive therapy (analysis utilising NMA of continuous data) and narrative exposure (analysis utilising NMA of dichotomous data) are the most cost-effective options at the NICE lower cost effectiveness threshold of £20,000/QALY. However, the probability of cognitive therapy being cost-effective is only 0.31. In the analysis utilising the NMA of dichotomous data, the probability of narrative exposure being cost-effective was above 0.70 at any cost effectiveness threshold.

Figure 186. Scenario D: Cost-effectiveness acceptability curves of interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); beneficial effect up to 3-month follow-up]



### Analysis utilising efficacy data from NMAs of dichotomous remission

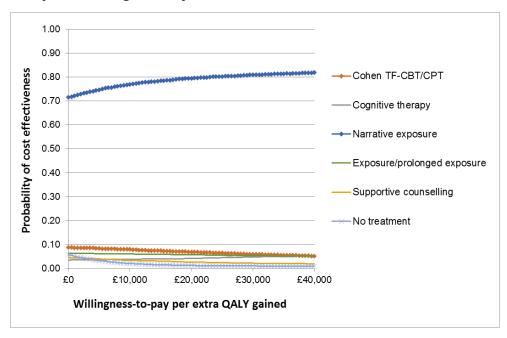
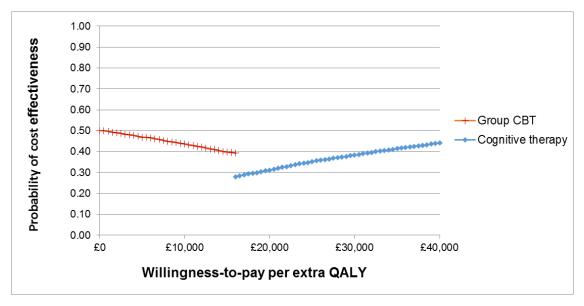
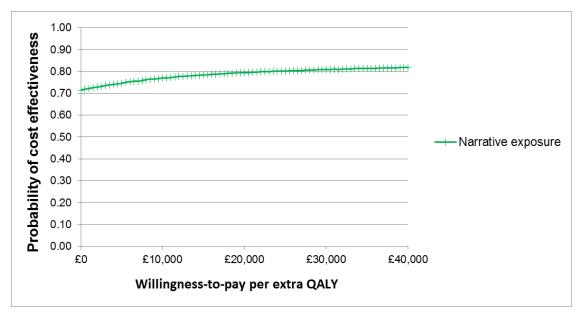


Figure 187 Scenario D: Cost-effectiveness acceptability frontier of interventions for the treatment of PTSD in children and young people [utility data from Shearer (2018); beneficial effect up to 3-month follow-up]



## Analysis utilising efficacy data from NMAs of dichotomous remission



Results were overall robust to changes in the annual risk of relapse.

## Discussion - conclusions, strengths and limitations of economic analysis

The guideline economic analysis assessed the cost effectiveness of a range of psychological interventions for the treatment of PTSD in children and young people. The interventions assessed were determined by the availability of efficacy data obtained from the NMAs that were conducted to inform this guideline. Interventions belonging to the TF-CBT class were assessed separately, as they differed in terms of related resource use and the results of the NMA suggested they had different efficacy as well. The base-case analysis conservatively assumed that the beneficial effect of interventions lasts only until treatment endpoint and that after this period, the probability of remission is equal to that of baseline treatment (no treatment). Alternative scenarios, using a beneficial effect of up to 3 months post-treatment (based on limited follow-up data) and a different set of utility values that translated into a narrower HRQoL benefit for people remitting were explored. The main analysis utilised continuous efficacy data, comprising changes in PTSD symptom scores, which were transformed to log-odds ratios of remission using a published formula. A secondary analysis utilised limited dichotomous efficacy data in an attempt to validate the conclusions of the main analysis. However, it needs to be noted that the definition of remission is different in the two analyses: in the analysis that derived remission from continuous data (changes in PTSD symptom scale scores), remission was defined as a final score below a hypothetical cut-off point on a PTSD symptom scale with an underlying normal distribution. In contrast, in the analysis that utilised dichotomous remission data, remission was defined as loss of PTSD diagnosis using DSM, ICD or similar criteria.

In the base-case analysis, the order of interventions from the most to the least cost-effective for the treatment of PTSD in children and young people was: cognitive therapy (TF-CBT), narrative exposure (TF-CBT), play therapy, prolonged exposure (TF-CBT), Cohen/CPT (TF-CBT), EMDR, parent training, group CBT (TF-CBT), family therapy, supportive counselling and no treatment. The probability of cognitive therapy being the most cost-effective treatment option was 0.78. It can be seen that with the exception of group CBT, all other interventions in the TF-CBT class are among the most cost-effective interventions of those assessed. In the secondary analysis that used dichotomous remission data, all interventions in the TF-CBT class were more cost-effective than no treatment; the order of interventions from the most to least cost-effective was: narrative exposure (TF-CBT), cognitive therapy (TF-CBT), prolonged exposure (TF-CBT), Cohen/CPT (TF-CBT), no treatment and supportive counselling. The probability of narrative exposure being the most cost-effective option was 0.62.

When a beneficial effect of up to 3 months post-treatment was assumed, the relative cost effectiveness of group CBT and Cohen/CPT (both TF-CBT) improved and the cost effectiveness of play therapy was reduced. The order of interventions became cognitive therapy (TF-CBT), Cohen/CPT (TF-CBT), group CBT (TF-CBT), narrative exposure (TF-CBT), parent training, prolonged exposure (TF-CBT), play therapy, EMDR, supportive counselling, family therapy, no treatment. The probability of cognitive therapy being the most cost-effective treatment option was 0.67. In the secondary analysis, the cost effectiveness of all interventions improved. Narrative exposure remained the most cost-effective intervention with a 0.81 probability, followed by cognitive therapy and then prolonged exposure.

When narrower utility benefits for remission and no beneficial effect beyond treatment endpoint were assumed, less costly interventions, such as EMDR and group CBT, were

favoured so that their relative cost effectiveness improved. The top-3 most cost-effective interventions remained the same with those of the base-case analysis and the order of interventions by cost effectiveness was as follows: cognitive therapy (TF-CBT), narrative exposure (TF-CBT), play therapy, group CBT (TF-CBT), EMDR, prolonged exposure (TF-CBT), parent training, Cohen/CPT (TF-CBT), family therapy, no treatment, supportive counselling. The probability of cognitive therapy being the most cost-effective treatment option was 0.59. In secondary analysis, only narrative exposure and cognitive therapy were more cost-effective than no treatment.

When narrower utility benefits for remission and a beneficial effect up to 3 months post-treatment were assumed, the order of interventions from most to least cost-effective became: cognitive therapy (TF-CBT), group CBT (TF-CBT), Cohen/CPT (TF-CBT), narrative exposure (TF-CBT), parent training, EMDR, play therapy, prolonged exposure (TF-CBT), family therapy, no treatment and supportive counselling. The probability of cognitive therapy being the most cost-effective intervention was only 0.31. In secondary analysis, the order of interventions by cost effectiveness was: narrative exposure, Cohen/CPT, prolonged exposure, cognitive therapy, supportive counselling, and no treatment. The probability of narrative exposure being the most cost-effective option was 0.79.

Results of the economic analysis were overall robust to the changes in the risk of relapse tested in deterministic sensitivity analysis.

Overall, individual forms of TF-CBT and, to a lesser degree, play therapy appear to be cost-effective in the treatment of children and young people with PTSD. Family therapy and supportive counselling do not appear to be cost-effective relative to other interventions and, under some scenarios, supportive counselling is less cost-effective than no treatment. In-between, there is another group of interventions (EMDR, group CBT and parent training) with modest relative cost effectiveness, which is affected by the alternative scenarios tested. The secondary analysis confirmed the cost effectiveness of individual forms of TF-CBT versus supportive counselling and no treatment, although the limited evidence did not allow further comparisons to be made.

The analysis utilised clinical effectiveness parameters derived from NMAs. This methodology enabled evidence synthesis from both direct and indirect comparisons between interventions, and allowed simultaneous inference on all treatments examined in pair-wise trial comparisons while respecting randomisation (Caldwell 2005; Lu & Ades, 2004). Global inconsistency checks indicated that there was no inconsistency between direct and indirect evidence considered in the 2 NMAs that utilised continuous data (PTSD changes in symptom scale scores). Regarding the NMA of dichotomous remission data, inconsistency checks were not relevant as there were no closed loops of direct evidence within the network.

The quality and limitations of RCTs considered in the NMAs have unavoidably impacted on the quality of the economic model clinical input parameters. For example, economic results may be have been affected by reporting and publication bias. Some interventions were informed by very limited evidence: cognitive therapy and parent training have been tested on 25 and 49 individuals, respectively, within the evidence base that informed the economic analysis. The evidence on dichotomous outcomes (remission) that informed the secondary analysis was even more limited; in particular, narrative exposure, cognitive therapy and

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prolonged exposure had been tested on 13, 26 and 50 people, respectively. However, the purpose of the secondary analysis was only to validate the conclusions reached using the results of the main analysis. It should be noted that, overall, the class of TF-CBT, in particular Cohen/CPT and group CBT within the class, had the most robust evidence base across all outcomes assessed in NMA.

The results of the NMAs of 1-4 month follow-up PTSD change score data and of the dichotomous remission data showed considerable uncertainty due to the small size of the included studies and the small total number of studies. Thus, results based on these data should be interpreted with caution. Nevertheless, the base-case economic analysis did not utilise the outputs of any of these NMAs. The NMA that informed the base-case economic analysis was based on more robust data and was characterised by moderate heterogeneity and no evidence of inconsistency.

The limitations characterising the data included in the NMAs and the NMA outputs informing the economic analyses should be considered when interpreting the cost effectiveness results.

The economic model did not consider discontinuation in the model structure due to the limited discontinuation data available. However, for the NMA that informed the economic analysis, ITT continuous data were extracted, where available. This means that discontinuation has been implicitly taken into account in the economic model outcomes. Moreover, the probabilistic analysis took into account the completion rates of the interventions assessed in the RCTs that informed the economic analysis, so that the number of sessions reflected, up to a degree, the attrition rates characterising each intervention.

The baseline risk of remission was estimated based on a large study that reported data on the course of PTSD derived from 1575 people with lifetime PTSD who had participated in 22 WHO World Mental Health surveys. Data on children and young people were possible to extract, so that remission data were directly relevant to the study population of the economic analysis. The risk of relapse was not possible to estimate using published evidence, and therefore was based on an assumption following advice from the committee. However, a range of values was tested in deterministic sensitivity analysis.

The time horizon of the analysis was 3 years, which were considered adequate to capture longer terms and costs associated with a course of treatment for PTSD without significant extrapolation over the course of PTSD.

Utility data used in the economic model were derived from a systematic review of studies reporting utility data for PTSD-related health states. The review included only two studies, each with different strengths and limitations. The economic analysis considered utility data from both studies in alternative scenarios.

Intervention costs were estimated based on relevant information provided in the studies included in the NMA supplemented by the committee's expert opinion, in order to reflect routine NHS practice. NHS and PSS costs incurred by children and young people with PTSD and those remitting from PTSD were taken from a small RCT due to lack of any other relevant evidence.

## Overall conclusions from the guideline economic analysis

Individual forms of TF-CBT and, to a lesser degree, play therapy appear to be cost-effective in the treatment of children and young people with PTSD. Family therapy and supportive counselling do not appear to be cost-effective relative to other interventions and, under some scenarios, supportive counselling is less cost-effective than no treatment. In-between, there is another group of interventions (EMDR, group CBT and parent training) with modest relative cost effectiveness. Results need to be interpreted with caution due to the limited evidence base characterising some of the interventions.

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Spiegelhalter DJ, Best NG, Carlin BP et al. (2002) Bayesian measures of model complexity and fit. Journal of the Royal Statistical Society: Series B 64(4), 583-616

van Valkenhoef G and Kuiper J (2016) gemtc: Network Meta-Analysis Using Bayesian Methods. R package. CRAN.

Yule W, Bolton D, Udwin O et al. (2000) The long-term psychological effects of a disaster experienced in adolescence: I: The incidence and course of PTSD. Journal of Child Psychology and Psychiatry 41(4), 503-11

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## **Appendix K – Excluded studies**

Excluded studies for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

#### **Clinical studies**

**Psychological: Trauma-focused CBT** 

| Study ID       | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|----------------|--|---|--|-------|
| Adelufosi 2017 | RQ 1.1-1.2 & 2.1-2.2 update              | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Adelufosi A, Edet B, Arikpo D, Aquaisua E, Meremikwu MM. Cognitive behavioral therapy for post-traumatic stress disorder, depression, or anxiety disorders in women and girls living with female genital mutilation: A systematic review. International Journal of Gynecology & Obstetrics. 2017 Feb 1;136(S1):56-9. |       |
| Capaldi 2016   | Cochrane allRQ update                    | Subgroup/secondary analysis of RCT already included   | Capaldi S, Asnaani A, Zandberg LJ, Carpenter JK, Foa EB. Therapeutic Alliance during Prolonged Exposure Versus Client-Centered Therapy for Adolescent Posttraumatic Stress Disorder. Journal of clinical psychology. 2016 Oct 1;72(10):1026-36.  |       |
| Cary 2012      | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-  | Cary CE, McMillen JC. The data behind the dissemination: A systematic review of trauma-  |       |

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| Study ID         | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|------------------|--|---|---|-------|
|                  |  | analysis results not appropriate to extract   | focused cognitive behavioral therapy for use with children and youth. Children and Youth Services Review. 2012 Apr 30;34(4):748-57.   |       |
| Chemtob 2008     | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided) | Chemtob CM, Luthra R. Effectiveness of Trauma-Focused Cognitive Behavioral Therapy in Treating Children With Post- Traumatic Stress Disorder [NCT00614068]. 2008. Available from: https://clinicaltrials.gov/ct2/show/ NCT00614068 [accessed 29.04.17]        |       |
| Cohen 2016       | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Intervention outside protocol   | Cohen JA, Mannarino AP, Jankowski K, Rosenberg S, Kodya S, Wolford GL. A randomized implementation study of trauma-focused cognitive behavioral therapy for adjudicated teens in residential treatment facilities. Child maltreatment. 2016 May;21(2):156-67. |       |
| Corcoran 2008    | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract     | Corcoran J, Pillai V. A meta-<br>analysis of parent-involved<br>treatment for child sexual abuse.<br>Research on Social Work<br>Practice. 2008 Sep;18(5):453-64.  |       |
| de Arellano 2014 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract     | de Arellano MA, Lyman DR,<br>Jobe-Shields L, George P,<br>Dougherty RH, Daniels AS,<br>Ghose SS, Huang L, Delphin-  |       |

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| Study ID       | Search                                   | Reason for exclusion   | Ref 1  | Ref 2 |
|----------------|--|--|--|-------|
|                |  |  | Rittmon ME. Trauma-focused cognitive-behavioral therapy for children and adolescents: Assessing the evidence. Psychiatric Services. 2014 May;65(5):591-602.  |       |
| Deblinger 1990 | 2004 GL (excluded)                       | Non-RCT (no control group)   | Deblinger E, McLEER SV, Henry D. Cognitive behavioral treatment for sexually abused children suffering post-traumatic stress: Preliminary findings. Journal of the American Academy of Child & Adolescent Psychiatry. 1990 Sep 1;29(5):747-52.   |       |
| Deblinger 2011 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Comparison outside protocol  | Deblinger E, Mannarino AP,<br>Cohen JA, Runyon MK, Steer<br>RA. Trauma-focused cognitive<br>behavioral therapy for children:<br>impact of the trauma narrative<br>and treatment length. Depression<br>and anxiety. 2011 Jan 1;28(1):67-<br>75.   |       |
| Dorsey 2014    | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Population not relevant for this review (to be considered for other relevant RQ) | Dorsey S, Pullmann MD, Berliner L, Koschmann E, McKay M, Deblinger E. Engaging foster parents in treatment: A randomized trial of supplementing Trauma-focused Cognitive Behavioral Therapy with evidence-based engagement strategies. Child abuse & neglect. 2014 Sep 30;38(9):1508-20. |       |
| Fernandez 2012 | Handsearch                               | Non-RCT (no control group)   | Fernandez, S., Cromer, L.D.,<br>Borntrager, C., Swopes*, R. &  |       |

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| Study ID                | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|-------------------------|--|---|---|-------|
|                         |  |   | Davis, J. L. (2012). A Case<br>Series: Cognitive-Behavioral<br>Treatment (Exposure, Relaxation,<br>and Rescripting Therapy) of<br>Trauma-Related Nightmares<br>Experienced by Children. Clinical<br>Case Studies, 12, 39-59.  |       |
| Forman-Hoffman<br>2013b | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Forman-Hoffman V, Knauer S, McKeeman J, Zolotor A, Blanco R, Lloyd S, et al. Child and adolescent exposure to trauma: comparative effectiveness of interventions addressing trauma other than maltreatment or family violence (Provisional abstract). Database of Abstracts of Reviews of Effects. 2013(2):1. |       |
| Gillies 2012            | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Gillies D, Taylor F, Gray C, O'Brien L, D'Abrew N. Psychological therapies for the treatment of post-traumatic stress disorder in children and adolescents. Cochrane Database of Systematic Reviews 2012, Issue 12. Art. No.: CD006726. DOI: 10.1002/14651858.CD006726.pu b2.                                 |       |
| Goenjian 1997           | Handsearch                               | Non-randomised group assignment   | Goenjian AK, Karayan I, Pynoos<br>RS, Minassian D, Najarian LM,<br>Steinberg AM, Fairbanks LA.<br>Outcome of psychotherapy<br>among early adolescents after<br>trauma. American Journal of  |       |

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| Study ID      | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|---------------|--|---|--|-------|
|               |  |   | Psychiatry. 1997 Apr 1;154(4):536-42.  |       |
| Haight 2012   | Handsearch                               | Intervention not targeted at PTSD symptoms  | Haight W, Black J, Sheridan K. A mental health intervention for rural, foster children from methamphetamine-involved families: Experimental assessment with qualitative elaboration. Children and youth services review. 2010 Oct 31;32(10):1446-57. |       |
| Harvey 2010   | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Harvey ST, Taylor JE. A meta-<br>analysis of the effects of<br>psychotherapy with sexually<br>abused children and adolescents.<br>Clinical Psychology Review. 2010<br>Jul 31;30(5):517-35.   |       |
| Hermenau 2013 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Population not relevant for this review (to be considered for other relevant RQ)                        | Hermenau, K., et al. (2013). Addressing post-traumatic stress and aggression by means of narrative exposure: A randomized controlled trial with ex-combatants in the eastern DRC. Journal of Aggression, Maltreatment and Trauma 22(8): 916-934.     |       |
| Hetrick 2010  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Hetrick SE, Purcell R, Garner B, Parslow R. Combined pharmacotherapy and psychological therapies for posttraumatic stress disorder (PTSD). Cochrane Database of Systematic Reviews 2010, Issue 7. Art. No.: CD007316. DOI:                           |       |

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| Study ID       | Search                                   | Reason for exclusion   | Ref 1   | Ref 2 |
|----------------|--|--|---|-------|
|                |  |  | 10.1002/14651858.CD007316.pu<br>b2.   |       |
| Holt 2014      | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Subgroup/secondary analysis of RCT already included  | Holt T, Jensen TK, Wentzel-<br>Larsen T. The change and the<br>mediating role of parental<br>emotional reactions and<br>depression in the treatment of<br>traumatized youth: results from a<br>randomized controlled study.<br>Child and adolescent psychiatry<br>and mental health. 2014 Apr<br>8;8(1):11. |       |
| Hyde 1995      | Handsearch                               | Intervention not targeted at PTSD symptoms   | Hyde C, Bentovim A, Monck E. Some clinical and methodological implications of a treatment outcome study of sexually abused children. Child Abuse & Neglect. 1995 Nov 1;19(11):1387-99.  |       |
| ISRCTN35018680 | Handsearch                               | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | ISRCTN35018680. A pilot randomised clinical trial of trauma-focused cognitive behaviour therapy for posttraumatic stress disorder (PTSD) in young children aged 3-8 years (PYCES). 2013. Available from: http://www.isrctn.com/ISRCTN35018680 [accessed 11.05.2017]   |       |
| ISRCTN58027256 | Handsearch                               | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | ISRCTN58027256. Identification and treatment within the Swedish Child and Adolescent Psychiatry Services of children exposed or subjected to intimate partner   |       |

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| Study ID          | Search                                   | Reason for exclusion   | Ref 1  | Ref 2 |
|-------------------|--|--|--|-------|
|                   |  |  | violence or child abuse: a randomised controlled trial. 2012. Available from: http://www.isrctn.com/ISRCTN58 027256 [accessed 11.05.2017]  |       |
| Jaberghaderi 2004 | 2004 GL (included)                       | Sample size (N<10/arm)   | Jaberghaderi, N., Greenwald, R., Rubin, A., Zand, S.O., Shiva Dolatabadi 1, S. (2004) A Comparison of CBT and EMDR for Sexually-abused Iranian Girls. Clinical Psychology and Psychotherapy 11, 358-368.   |       |
| Kalantari 2012    | Handsearch                               | Population outside scope: Trials of people with traumatic grief  | Kalantari M, Yule W, Dyregrov A, Neshatdoost H, Ahmadi SJ. Efficacy of writing for recovery on traumatic grief symptoms of Afghani refugee bereaved adolescents: A randomized control trial. OMEGA-Journal of death and dying. 2012 Oct;65(2):139-50.  |       |
| Kameoka 2013      | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | Kameoka S. Randomized controlled trial on the efficacy of the Trauma-Focused Cognitive Behavioral Therapy for children with posttraumatic stress disorder [JPRN-UMIN000010699]. Available from: https://upload.umin.ac.jp/cgiopen-bin/ctr_e/ctr_view.cgi?recptno=R 000012501 [accessed 30.04.17] |       |
| Kane 2016         | RQ 1.1-1.2 & 2.1-2.2 update              | Subgroup/secondary analysis that is not relevant   | Kane JC, Murray LK, Cohen J,<br>Dorsey S, Skavenski van Wyk S,   |       |

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| Study ID      | Search                                   | Reason for exclusion   | Ref 1  | Ref 2 |
|---------------|--|--|--|-------|
|               |  |  | Galloway Henderson J, Imasiku M, Mayeya J, Bolton P. Moderators of treatment response to trauma-focused cognitive behavioral therapy among youth in Zambia. Journal of Child Psychology and Psychiatry. 2016 Oct 1;57(10):1194-202.  |       |
| Kenardy 2012  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | Kenardy J. Comparison of cognitive-behavioural treatments for children with post-traumatic stress disorder (PTSD) following an accidental injury: a multicentre randomised controlled trial [ISRCTN79049138]. 2012. Available from: http://www.isrctn.com/ISRCTN79049138 [accessed 30.04.17] |       |
| Kowalik 2011  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract          | Kowalik J, Weller J, Venter J, Drachman D. Cognitive behavioral therapy for the treatment of pediatric posttraumatic stress disorder: A review and meta-analysis. Journal of Behavior Therapy and Experimental Psychiatry. 2011 Sep 30;42(3):405-13.   |       |
| Leenarts 2013 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract          | Leenarts LE, Diehle J, Doreleijers TA, Jansma EP, Lindauer RJ. Evidence-based treatments for children with trauma-related psychopathology as a result of childhood maltreatment: a   |       |

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| Study ID          | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|-------------------|--|---|--|-------|
|                   |  |   | systematic review. European child & adolescent psychiatry. 2013 May 1;22(5):269-83.  |       |
| Lenz 2015         | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract     | Lenz AS, Hollenbaugh KM. Meta-<br>analysis of trauma-focused<br>cognitive behavioral therapy for<br>treating PTSD and co-occurring<br>depression among children and<br>adolescents. Counseling<br>Outcome Research and<br>Evaluation. 2015 Jun;6(1):18-32. |       |
| McLean 2015b      | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Subgroup/secondary analysis of RCT already included   | McLean CP, Su YJ, Foa EB. Mechanisms of symptom reduction in a combined treatment for comorbid posttraumatic stress disorder and alcohol dependence. Journal of consulting and clinical psychology. 2015 Jun;83(3):655.                                    |       |
| McLean 2017       | RQ 1.1-1.2 & 2.1-2.2 update              | Subgroup/secondary analysis of RCT already included   | McLean CP, Su YJ, Carpenter JK, Foa EB. Changes in PTSD and depression during prolonged exposure and client-centered therapy for PTSD in adolescents. Journal of Clinical Child & Adolescent Psychology. 2017 Jul 4;46(4):500-10.                          |       |
| Miller-Graff 2016 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Miller-Graff LE, Campion K. Interventions for posttraumatic stress with children exposed to violence: factors associated with treatment success. Journal of clinical psychology. 2015 Nov 1.   |       |

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| Study ID     | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|--------------|--|---|---|-------|
| Morina 2016  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract         | Morina N, Koerssen R, Pollet TV. Interventions for children and adolescents with posttraumatic stress disorder: A meta-analysis of comparative outcome studies. Clinical Psychology Review. 2016 Jul 31;47:41-54.   |       |
| Morina 2017b | RQ 1.1-1.2 & 2.1-2.2 update              | Systematic review with no new useable data and any meta-analysis results not appropriate to extract         | Morina N, Malek M, Nickerson A, Bryant RA. Psychological interventions for post-traumatic stress disorder and depression in young survivors of mass violence in low-and middle-income countries: meta-analysis. The British Journal of Psychiatry. 2017 Apr 1;210(4):247-54.  |       |
| Murray 2015  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Efficacy or safety data cannot be extracted   | Murray LK, Skavenski S, Kane JC, Mayeya J, Dorsey S, Cohen JA, Michalopoulos LT, Imasiku M, Bolton PA. Effectiveness of Trauma-Focused Cognitive Behavioral Therapy Among Trauma-Affected Children in Lusaka, Zambia: A Randomized Clinical Trial. JAMA Pediatr. 2015 Aug;169(8):761-9. doi: 10.1001/jamapediatrics.2015.058 0. |       |
| NCT00073684  | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided) | NCT00073684. Young Sexually<br>Abused Children: Optimal CBT<br>Strategies. 2003. Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT00073684 [accessed<br>11.05.2017]  |       |

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| Study ID    | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|-------------|--|---|--|-------|
| NCT00893750 | Handsearch                               | Dissertation  | NCT00893750. Effects of<br>Trauma-Therapy and Truth<br>Education, Conflict Resolution<br>and Social Skills Trainings and<br>Traditional Ways of Coping in<br>Northern Uganda. 2009.<br>Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT00893750 [accessed<br>11.05.17] |       |
| NCT02334566 | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided) | NCT02334566. Lending a Hand to Our Future: Documenting, Assessing and Treating Posttraumatic Stress Disorder in Refugee Children and Youth. 2014. Available from: https://clinicaltrials.gov/ct2/show/NCT02334566 [accessed 11.05.2017]  |       |
| NCT02402205 | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided) | NCT02402205. TF-CBT for<br>Adjudicated Youth in Residential<br>Treatment. 2015. Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT02402205 [accessed<br>11.05.2017]  |       |
| Nenova 2013 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract     | Nenova M, Morris L, Paul L, Li Y,<br>Applebaum A, DuHamel K.<br>Psychosocial interventions with<br>cognitive-behavioral components<br>for the treatment of cancer-<br>related traumatic stress<br>symptoms: a review of<br>randomized controlled trials. J                           |       |

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| Study ID         | Search   | Reason for exclusion  | Ref 1   | Ref 2   |
|------------------|--|---|---|---|
|                  |  |   | Cogn Psychother. 2013 Jan 1;27(3):258-84.   |   |
| Nixon 2012a/2017 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) AND RQ 1.1-1.2 & 2.1-2.2 update | Comparison outside protocol   | Nixon RD, Sterk J, Pearce A. A Randomized Trial of Cognitive Behavior Therapy and Cognitive Therapy for Children with Posttraumatic Stress Disorder Following Single-Incident Trauma. Journal of Abnormal Child Psychology. 2012 Apr 1;40(3):327. | Nixon RD, Sterk J, Pearce A, Weber N. A randomized trial of cognitive behavior therapy and cognitive therapy for children with posttraumatic stress disorder following single-incident trauma: Predictors and outcome at 1-year follow-up. Psychological Trauma: Theory, Research, Practice, and Policy. 2017 Jul;9(4):471. |
| Ormaugh 2014     | RQ 1.1-1.2 & 2.1-2.2 (searches combined)                                 | Subgroup/secondary analysis of RCT already included   | Ormhaug SM, Jensen TK, Wentzel-Larsen T, Shirk SR. The therapeutic alliance in treatment of traumatized youths: Relation to outcome in a randomized clinical trial. Journal of consulting and clinical psychology. 2014 Feb;82(1):52.             |   |
| Parsons 2008     | RQ 1.1-1.2 & 2.1-2.2 (searches combined)                                 | Systematic review with no new useable data and any meta-analysis results not appropriate to extract     | Parsons TD, Rizzo AA. Affective outcomes of virtual reality exposure therapy for anxiety and specific phobias: A meta-analysis. Journal of behavior therapy and experimental psychiatry. 2008 Sep 30;39(3):250-61.                                |   |
| Reynolds 2012    | RQ 1.1-1.2 & 2.1-2.2 (searches combined)                                 | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Reynolds S, Wilson C, Austin J,<br>Hooper L. Effects of<br>psychotherapy for anxiety in<br>children and adolescents: A<br>meta-analytic review. Clinical  |   |

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| Study ID                      | Search                                   | Reason for exclusion  | Ref 1   | Ref 2  |
|-------------------------------|--|---|---|--|
| -                             |  |   | psychology review. 2012 Jun 30;32(4):251-62.  |  |
| Rolfsnes 2011                 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Rolfsnes ES, Idsoe T. School-<br>based intervention programs for<br>PTSD symptoms: A review and<br>meta-analysis. Journal of<br>Traumatic Stress. 2011 Apr<br>1;24(2):155-65.   |  |
| Salloum 2008                  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Comparison outside protocol   | Salloum A, Overstreet S. Evaluation of individual and group grief and trauma interventions for children post disaster. Journal of Clinical Child & Adolescent Psychology. 2008 Jul 14;37(3):495-507.  |  |
| Salloum 2014                  | Handsearch                               | Sample size (N<10/arm)  | Salloum A, Robst J, Scheeringa MS, Cohen JA, Wang W, Murphy TK, Tolin DF, Storch EA. Step one within stepped care traumafocused cognitive behavioral therapy for young children: a pilot study. Child Psychiatry Hum Dev. 2014 Feb;45(1):65-77. doi: 10.1007/s10578-013-0378-6. |  |
| Salloum 2015                  | Handsearch                               | Sample size (N<10/arm)  | Salloum A, Small BJ, Robst J,<br>Scheeringa MS, Cohen JA,<br>Storch EA. Stepped and standard<br>care for childhood trauma: A pilot<br>randomized clinical trial.<br>Research on Social Work<br>Practice. 2015 Sep<br>24:1049731515601898.                                       | Salloum A, Scheeringa MS,<br>Cohen JA, Storch EA.<br>Responder Status Criterion for<br>Stepped Care Trauma-<br>Focused Cognitive Behavioral<br>Therapy for Young Children.<br>Child Youth Care Forum. 2015<br>Feb;44(1):59-78. |
| Scheeringa<br>2011/Weems 2013 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Non-randomised group assignment   | Scheeringa MS, Weems CF,<br>Cohen JA, Amaya-Jackson L,  | Weems CF, Scheeringa MS. Maternal depression and   |

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| Study ID       | Search                                   | Reason for exclusion   | Ref 1   | Ref 2  |
|----------------|--|--|---|--|
|                |  |  | Guthrie D. Trauma-focused cognitive-behavioral therapy for posttraumatic stress disorder in three-through six year-old children: A randomized clinical trial. Journal of Child Psychology and Psychiatry. 2011 Aug 1;52(8):853-60.  | treatment gains following a cognitive behavioral intervention for posttraumatic stress in preschool children. Journal of anxiety disorders. 2013 Jan 31;27(1):140-6. |
| Scott 2005     | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract          | Scott RW, Mughelli K, Deas D.<br>An overview of controlled studies<br>of anxiety disorders treatment in<br>children and adolescents. Journal<br>of the National Medical<br>Association. 2005 Jan;97(1):13.  |  |
| Silverman 2008 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract              | Silverman WK, Ortiz CD,<br>Viswesvaran C, Burns BJ, Kolko<br>DJ, Putnam FW, Amaya-Jackson<br>L. Evidence-based psychosocial<br>treatments for children and<br>adolescents exposed to traumatic<br>events. Journal of Clinical Child &<br>Adolescent Psychology. 2008<br>Mar 3;37(1):156-83. |  |
| Stallard 2006b | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | Stallard P. A pilot randomised trial to determine the efficacy of early cognitive behaviour therapy (CBT) versus delayed treatment for children with significant post-traumatic reactions [ISRCTN05595708]. 2006. Available from: http://www.isrctn.com/ISRCTN05595708 [accessed 30.04.17]  |  |

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| Study ID      | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|---------------|--|---|--|-------|
| Swain 2013    | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Swain J, Hancock K, Hainsworth C, Bowman J. Acceptance and commitment therapy in the treatment of anxiety: a systematic review. Clinical psychology review. 2013 Dec 31;33(8):965-78.  |       |
| Taylor 2004   | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Taylor TL, Chemtob CM. Efficacy of treatment for child and adolescent traumatic stress. Archives of pediatrics & adolescent medicine. 2004 Aug 1;158(8):786-91.  |       |
| Townsend 2008 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Protocol  | Townsend E, Walker DM,<br>Sargeant S, Stocker O, Vostanis<br>P, Sithole J, Hawton KKE.<br>Interventions for mood and<br>anxiety disorders, and self harm<br>in young offenders. Cochrane<br>Database of Systematic Reviews<br>2008, Issue 2. Art. No.:<br>CD007195. DOI:<br>10.1002/14651858.CD007195. |       |
| Trask 2011    | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-<br>analysis results not appropriate to extract | Trask EV, Walsh K, DiLillo D. Treatment effects for common outcomes of child sexual abuse: A current meta-analysis. Aggression and violent behavior. 2011 Feb 28;16(1):6-19.   |       |
| Tutus 2017    | RQ 1.1-1.2 & 2.1-2.2 update              | Efficacy or safety data cannot be extracted   | Tutus D, Pfeiffer E, Rosner R,<br>Sachser C, Goldbeck L.<br>Sustainability of Treatment<br>Effects of Trauma-focused<br>Cognitive-behavioral Therapy for   |       |

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| Study ID      | Search     | Reason for exclusion   | Ref 1  | Ref 2 |
|---------------|------------|--|--|-------|
|               |            |  | Children and Adolescents:<br>Findings from 6-and 12-month<br>Follow-ups. Psychotherapy and<br>psychosomatics. 2017;86(6):379-<br>81.   |       |
| UMIN000010699 | Handsearch | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | Randomized controlled trial on the efficacy of the Trauma-Focused Cognitive Behavioral Therapy for children with posttraumatic stress disorder, https://upload.umin.ac.jp/cgiopen-bin/ctr_e/ctr_view.cgi?recptno=R 000012501 |       |

Psychological: Non-trauma-focused CBT

| Study ID      | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|---------------|--|---|---|-------|
| James<br>2015 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | James AC, James G, Cowdrey FA, Soler A, Choke A. Cognitive behavioural therapy for anxiety disorders in children and adolescents. Cochrane Database of Systematic Reviews 2015, Issue 2. Art.No.: CD004690. DOI: 10.1002/14651858.CD004690.pu b4.     |       |
| March<br>1998 | 2004 GL (excluded)                       | Non-randomised group assignment   | March, J. S., Amaya-Jackson, L., Murray, M. C., & Schulte, A. (1998). Cognitive-behavioral psychotherapy for children and adolescents with posttraumatic stress disorder after a single-incident stressor. Journal of the American Academy of Child & |       |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Study ID          | Search                                   | Reason for exclusion                       | Ref 1   | Ref 2 |
|-------------------|--|--|---|-------|
|                   |  |  | Adolescent Psychiatry, 37, 585-593.   |       |
| Mitchell<br>2011  | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Intervention not targeted at PTSD symptoms | Mitchell P, Smedley K, Kenning C, McKee A, Woods D, Rennie CE, Bell RV, Aryamanesh M, Dolan M. Cognitive behaviour therapy for adolescent offenders with mental health problems in custody. Journal of adolescence. 2011 Jun 30;34(3):433-43.   |       |
| Schaeffer<br>2013 | Handsearch                               | Intervention not targeted at PTSD symptoms | Schaeffer, C., Swenson, C.,<br>Tuerk, E. and Henggler, S.<br>(2013) Comprehensive treatment<br>for co-occurring child<br>maltreatment and parental<br>substance abuse: Outcomes from<br>a 24-month pilot study of the<br>MST-Building Stronger Families<br>program, Child Abuse and<br>Neglect, 37, 596-607 |       |

Psychological: Behavioural therapy

| Study ID         | Search             | Reason for exclusion  | Ref 1   | Ref 2 |
|------------------|--------------------|---|---|-------|
| Berliner<br>1996 | 2004 GL (excluded) | Intervention not targeted at PTSD symptoms  | Berliner L, Saunders BE. Treating fear and anxiety in sexually abused children: Results of a controlled 2-year follow-up study. Child maltreatment. 1996 Nov 1;1(4):294-309 |       |
| Lustig<br>2008   | Handsearch         | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Lustig, S., Tennakoon, L. (2008)<br>Testimonials, narratives, stories<br>and drawings: child refugees as<br>witnesses, Child and Adolescent                                 |       |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

| Study ID            | Search     | Reason for exclusion | Ref 1   | Ref 2   |
|---------------------|------------|----------------------|---|---|
|                     |            |                      | Psychiatric Clinics of North<br>America, 17, 569-584                                  |   |
| Macfarla<br>ne 1986 | Handsearch | Book Section         | MacFarlane K, Cunningham C (1986), Steps to H~althy Touching. Mt Dora, FL: Kidsrights | Cohen, J., Bernet, W., Dunne, J., Adair, M., Arnold, V., Benson, R., Bukstein, O., Kinlan, J., McClellan, J., Rue, D. & Sloan, E. (1998) Practice parameters for the assessment and treatment of children and adolescents with posttraumatic stress disorder, Journal of the American Academy of Child and Adolescent Psychiatry, 37, |

Psychological: Psychologically-focused debriefing

| Study ID       | Search                                   | Reason for exclusion            | Ref 1  | Ref 2   |
|----------------|--|---------------------------------|--|---|
| Pynoos<br>1988 | Handsearch                               | Commentary                      | Pyno os RS, Nader K (1988),<br>Psychological first aid and<br>treatment approach to child ren<br>exposed to community violence:<br>research implications, Trauma<br>Stress 1:445 - 473 | Cohen, J., Bernet, W., Dunne, J., Adair, M., Arnold, V., Benson, R., Bukstein, O., Kinlan, J., McClellan, J., Rue, D. & Sloan, E. (1998) Practice parameters for the assessment and treatment of children and adolescents with posttraumatic stress disorder, Journal of the American Academy of Child and Adolescent Psychiatry, 37, |
| Thabet<br>2005 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Non-randomised group assignment | Thabet AA, Vostanis P, Karim K. Group crisis intervention for children during ongoing war conflict. European Child & Adolescent Psychiatry. 2005 Aug 1;14(5):262-9.                    |   |

Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

Psychological: Eye movement desensitisation and reprocessing (EMDR)

| Study ID                              | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|---------------------------------------|--|---|---|-------|
| Field<br>2011                         | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Field A, Cottrell D. Eye movement desensitization and reprocessing as a therapeutic intervention for traumatized children and adolescents: a systematic review of the evidence for family therapists. Journal of Family Therapy. 2011 Nov 1;33(4):374-88.   |       |
| Greyber<br>2012                       | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Greyber LR, Dulmus CN, Cristalli ME. Eye movement desensitization reprocessing, posttraumatic stress disorder, and trauma: A review of randomized controlled trials with children and adolescents. Child and Adolescent Social Work Journal. 2012 Oct 1;29(5):409-25.   |       |
| Hassanz<br>adeh<br>Moghadd<br>am 2016 | Handsearch                               | Intervention not targeted at PTSD symptoms  | Hassanzadeh Moghaddam M,<br>Khalatbari J. Investigating the<br>Effectiveness of Eye Movement<br>Desensitization and<br>Reprocessing (EMDR) on<br>Children with Post-Traumatic<br>Stress Disorder (Traffic<br>Accident). The International<br>Journal of Indian Psychology,<br>Volume 3, Issue 3, No. 11. 2016<br>Jun 29:45. |       |
| Kemp<br>2010                          | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Efficacy or safety data cannot be extracted   | Kemp M, Drummond P,<br>McDermott B. A wait-list<br>controlled pilot study of eye  |       |

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| Study ID           | Search                                   | Reason for exclusion   | Ref 1   | Ref 2 |
|--------------------|--|--|---|-------|
|                    |  |  | movement desensitization and reprocessing (EMDR) for children with post-traumatic stress disorder (PTSD) symptoms from motor vehicle accidents. Clinical child psychology and psychiatry. 2010 Jan 1;15(1):5-25.  |       |
| Rodenbu<br>rg 2009 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract              | Rodenburg R, Benjamin A, de<br>Roos C, Meijer AM, Stams GJ.<br>Efficacy of EMDR in children: A<br>meta-analysis. Clinical<br>Psychology Review. 2009 Nov<br>30;29(7):599-606.   |       |
| Roos<br>2013       | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Unpublished (registered on clinical trials registry and author contacted for full trial report but not provided) | Roos C. A Randomized Comparison of Eye Movement Desensitization and Reprocessing (EMDR) and Cognitive Behavioral Writing Therapy (CBWT) in pediatric posttraumatic stress disorder following single- incident trauma [NTR3870]. 2013. Available from: http://www.trialregister.nl/trialreg/ admin/rctview.asp?TC=3870 [accessed 30.04.17] |       |
| Rubin<br>2001      | Handsearch                               | Intervention not targeted at PTSD symptoms   | Rubin A, Bischofshausen S, Conroy-Moore K, Dennis B, Hastie M, Melnick L, Reeves D, Smith T. The effectiveness of EMDR in a child guidance center. Research on Social Work Practice. 2001 Jul;11(4):435-57.   |       |

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| Study ID        | Search                      | Reason for exclusion  | Ref 1   | Ref 2 |
|-----------------|-----------------------------|---|---|-------|
| Verardo<br>2017 | RQ 1.1-1.2 & 2.1-2.2 update | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Verardo AR, Cioccolanti E. TRAUMATIC EXPERIENCES AND EMDR IN CHILDHOOD AND ADOLESCENCE. A REVIEW OF THE SCIENTIFIC LITERATURE ON EFFICACY STUDIES. Clinical Neuropsychiatry. 2017 Oct 1(5). |       |

**Psychological: Hypnotherapy** 

| Study ID        | Search                                   | Reason for exclusion          | Ref 1   | Ref 2 |
|-----------------|--|-------------------------------|---|-------|
| Lesmana<br>2009 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Intervention outside protocol | Lesmana CB, Suryani LK,<br>Jensen GD, Tiliopoulos N. A<br>spiritual-hypnosis assisted<br>treatment of children with PTSD<br>after the 2002 Bali terrorist<br>attack. American Journal of<br>Clinical Hypnosis. 2009 Jul<br>1;52(1):23-34. |       |

Psychological: Psychodynamic therapies

| Study ID            | Search     | Reason for exclusion       | Ref 1  | Ref 2  |
|---------------------|------------|----------------------------|--|--|
| Gaensba<br>uer 1994 | Handsearch | Non-RCT (no control group) | Gaensbauer TJ (1994) . Therapeutic work with a traumatized toddler. Psycboanal StudyChild 49:412-433 | Cohen, J., Bernet, W., Dunne, J., Adair, M., Arnold, V., Benson, R., Bukstein, O., Kinlan, J., McClellan, J., Rue, D. & Sloan, E. (1998) Practice parameters for the assessment and treatment of children and adolescents with posttraumatic stress disorder, Journal of the |

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| Study ID        | Search             | Reason for exclusion        | Ref 1  | Ref 2  |
|-----------------|--------------------|-----------------------------|--|--|
|                 |                    |                             |  | American Academy of Child and Adolescent Psychiatry, 37, |
| Trowell<br>2002 | 2004 GL (included) | Comparison outside protocol | Trowell, J., Kolvin, I., Weeramanthri, T., Sadowski, H., Berelowitz, M., Glaser, D. et al. (2002). Psychotherapy for sexually abused girls: psychopathological outcome findings and patterns of change. Br.J Psychiatry, 180, 234-247. |  |

#### **Psychological: Psychoeducation**

| Study ID               | Search                                   | Reason for exclusion  | Ref 1   | Ref 2 |
|------------------------|--|---|---|-------|
| Adler-<br>Nevo<br>2005 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Adler-Nevo G, Manassis K. Psychosocial treatment of pediatric posttraumatic stress disorder: the neglected field of single-incident trauma. Depression and Anxiety. 2005 Jan 1;22(4):177-89.  |       |
| Ager 2011              | Handsearch                               | Outcome measures are not validated  | Ager A, Akesson B, Stark L, Flouri E, Okot B, McCollister F, Boothby N. The impact of the school-based Psychosocial Structured Activities (PSSA) program on conflict-affected children in northern Uganda. Journal of Child Psychology and Psychiatry. 2011 Nov 1;52(11):1124-33. |       |
| Kazdin<br>2002         | 2004 GL (excluded)                       | Commentary  | Kazdin A.(2002) Comment on a school based psychosocial intervention was effective in  |       |

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| Study ID            | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|---------------------|--|---|--|-------|
|                     |  |   | children with persistent post-<br>disaster trauma symptoms.) Evid<br>Based Ment Health. 2002<br>Aug;5(3):76.   |       |
| NCT0075<br>1946     | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided) | NCT00751946. Girls In Recovery<br>From Life Stress (GIRLS) Study.<br>2008. Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT00751946 [accessed<br>11.05.2017]   |       |
| Peltonen<br>2012    | Handsearch                               | Non-randomised group assignment   | Peltonen K, Qouta S, El Sarraj E, Punamäki RL. Effectiveness of school-based intervention in enhancing mental health and social functioning among waraffected children. Traumatology. 2012 Dec;18(4):37-46.  |       |
| Salloum<br>2012     | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Comparison outside protocol   | Salloum A, Overstreet S. Grief and trauma intervention for children after disaster: Exploring coping skills versus trauma narration. Behaviour research and therapy. 2012 Mar 31;50(3):169-79.   |       |
| Santacro<br>ce 2010 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Sample size (N<10/arm)  | Judge Santacroce S, Asmus K, Kadan-Lottick N, Grey M. Feasibility and preliminary outcomes from a pilot study of coping skills training for adolescent—Young adult survivors of childhood cancer and their parents. Journal of pediatric oncology nursing. 2010 Jan;27(1):10-20. |       |

**Psychological: Counselling** 

| Study ID                 | Search     | Reason for exclusion | Ref 1   | Ref 2   |
|--------------------------|------------|----------------------|---|---|
| Lowenste in 1995         | Handsearch | Commentary           | Lowenstein LB (1995), The resolution scrapbook as an aid in the treatment of traumatized children. Child ~/far~ 74:899- 904   | Cohen, J., Bernet, W., Dunne, J., Adair, M., Arnold, V., Benson, R., Bukstein, O., Kinlan, J., McClellan, J., Rue, D. & Sloan, E. (1998) Practice parameters for the assessment and treatment of children and adolescents with posttraumatic stress disorder, Journal of the American Academy of Child and Adolescent Psychiatry, 37, |
| Schauer<br>2005/201<br>1 | Handsearch | Book Section         | Schauer M, Neuner F, Elbert T<br>(2005/2011): Narrative Exposure<br>Therapy. A Short-Term<br>Intervention for Traumatic Stress<br>Disorders. 2nd Ed. Cambridge/<br>Göttingen: Hogrefe & Huber<br>Publishers |   |
| Sullivan<br>1994         | Handsearch | Commentary           | Sullivan JM, Evans K (1994),<br>Integrated treatment for the<br>survivor of childhood trauma who<br>is chemically dependent. }<br>Psycboactiue Drugs 26:369-378   | Cohen, J., Bernet, W., Dunne, J., Adair, M., Arnold, V., Benson, R., Bukstein, O., Kinlan, J., McClellan, J., Rue, D. & Sloan, E. (1998) Practice parameters for the assessment and treatment of children and adolescents with posttraumatic stress disorder, Journal of the American Academy of Child and Adolescent Psychiatry, 37, |

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Psychological: Self-help (without support)

| Study ID        | Search                                   | Reason for exclusion  | Ref 1  | Ref 2 |
|-----------------|--|---|--|-------|
| Pennant<br>2015 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract | Pennant ME, Loucas CE, Whittington C, Creswell C, Fonagy P, Fuggle P, Kelvin R, Naqvi S, Stockton S, Kendall T, Group EA. Computerised therapies for anxiety and depression in children and young people: A systematic review and meta-analysis. Behaviour research and therapy. 2015 Apr 30;67:1-8. |       |

**Psychological: Parent training/family interventions** 

| Study        | D Search                                 | Reason for exclusion                        | Ref 1   | Ref 2 |
|--------------|--|---|---|-------|
| Saxe<br>2012 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Efficacy or safety data cannot be extracted | Saxe, G. N., Heidi Ellis, B.,<br>Fogler, J., Navalta, C. P. (2012)<br>Innovations in Practice:<br>Preliminary evidence for effective<br>family engagement in treatment<br>for child traumatic stress-trauma<br>systems therapy approach to<br>preventing dropout, Child and<br>Adolescent Mental Health, 17,<br>58-61 |       |

**Psychosocial: Art therapy** 

| Study ID        | Search                                   | Reason for exclusion            | Ref 1  | Ref 2 |
|-----------------|--|---------------------------------|--|-------|
| Brillantes<br>- | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Non-randomised group assignment | Brillantes-Evangelista G. An evaluation of visual arts and poetry as therapeutic |       |

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| Study ID             | Search                                   | Reason for exclusion                        | Ref 1   | Ref 2 |
|----------------------|--|---|---|-------|
| Evangelis<br>ta 2013 |  |   | interventions with abused adolescents. The Arts in Psychotherapy. 2013 Feb 28;40(1):71-84.  |       |
| Raider<br>2008       | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Sample size (N<10/arm)                      | Raider MC, Steele W, Delillo-<br>Storey M, Jacobs J, Kuban C.<br>Structured sensory therapy<br>(SITCAP-ART) for traumatized<br>adjudicated adolescents in<br>residential treatment. Residential<br>Treatment for Children & Youth.<br>2008 Sep 4;25(2):167-85.  |       |
| Schreier<br>2005     | Handsearch                               | Efficacy or safety data cannot be extracted | Schreier H, Ladakakos C, Morabito D, Chapman L, Knudson MM. Posttraumatic stress symptoms in children after mild to moderate pediatric trauma: a longitudinal examination of symptom prevalence, correlates, and parent-child symptom reporting. Journal of Trauma and Acute Care Surgery. 2005 Feb 1;58(2):353-63. |       |

#### **Psychosocial: Music therapy**

| Study ID      | Search     | Reason for exclusion            | Ref 1  | Ref 2 |
|---------------|------------|---------------------------------|--|-------|
| Baker<br>2006 | Handsearch | Non-randomised group assignment | Baker F, Jones C. The effect of music therapy services on classroom behaviours of newly arrived refugee students in Australia—a pilot study. Emotional and Behavioural |       |

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| Study ID Search | Reason for exclusion | Ref 1                                  | Ref 2 |
|-----------------|----------------------|--|-------|
|                 |                      | Difficulties. 2006 Dec 1;11(4):249-60. |       |

#### **Psychosocial: Meditation**

| <i>-</i> |                   |  |  |   |       |
|----------|-------------------|--|--|---|-------|
|          | Study ID          | Search                                   | Reason for exclusion   | Ref 1   | Ref 2 |
|          | Hartman<br>n 2012 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Systematic review with no new useable data and any meta-analysis results not appropriate to extract                              | Hartmann F, Vlieger AM. Effects of mind–body therapies in children. Focus on Alternative and Complementary Therapies. 2012 Jun 1;17(2):91-6.  |       |
|          | NCT0020<br>2709   | Handsearch                               | Population outside scope: <80% of the study's participants are eligible for the review and disaggregated data cannot be obtained | NCT00202709. Can Thought<br>Field Therapy (TFT) be Helpful<br>for Patients With an Anxiety<br>Disorder, a Prospective,<br>Randomized Pilot Study With<br>Wait List as Control Group.<br>Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT00202709 [accessed<br>14/06/17] |       |
|          | NCT0159<br>5477   | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided)                      | NCT01595477. A Randomized<br>Controlled Study of Mind-Body<br>Skills Groups for the Treatment of<br>War-Related Trauma in Children<br>in Gaza. 2012. Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT01595477 [accessed<br>11.05.2017]                                  |       |
|          | NCT0159<br>5490   | Handsearch                               | Unpublished (registered on clinical trials.gov and author contacted for full trial report but not provided)                      | NCT01595490. A Randomized<br>Controlled Study of Mind-Body<br>Skills Groups for the Treatment of  |       |

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| Study ID | Search | Reason for exclusion | Ref 1  | Ref 2 |
|----------|--------|----------------------|--|-------|
|          |        |                      | War-Related Trauma in<br>Adolescents in Gaza. 2012.<br>Available from:<br>https://clinicaltrials.gov/ct2/show/<br>NCT01595490 [accessed<br>11.05.2017] |       |

**Psychosocial: Peer support** 

| Study I         | D Search           | Reason for exclusion                        | Ref 1  | Ref 2 |
|-----------------|--------------------|---|--|-------|
| Fantuzz<br>1996 | zo Handsearch      | Intervention not targeted at PTSD symptoms  | Fantuzzo J, Sutton-Smith B,<br>Atkins M, Meyers R, Stevenson<br>H, Coolahan K, Weiss A, Manz P.<br>Community-based resilient peer<br>treatment of withdrawn<br>maltreated preschool children.<br>Journal of Consulting and Clinical<br>Psychology. 1996<br>Dec;64(6):1377. |       |
| Fantuzz<br>2005 | zo Handsearch      | Intervention not targeted at PTSD symptoms  | Fantuzzo J, Manz P, Atkins M, Meyers R. Peer-mediated treatment of socially withdrawn maltreated preschool children: Cultivating natural community resources. Journal of Clinical Child and Adolescent Psychology. 2005 May 1;34(2):320-5.                                 |       |
| Hardin<br>2002  | 2004 GL (excluded) | Efficacy or safety data cannot be extracted | Hardin, S.B. Weinrich, S.;<br>Weinrich, M.; Garrison, C.; Addy,<br>C. & Hardin, T.L. (2002) Effects<br>of long-term psychological<br>nursing intervention on<br>adolescents exposed to<br>catastrophic stress. Issues in   |       |

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| Study ID           | Search                                   | Reason for exclusion                        | Ref 1  | Ref 2 |
|--------------------|--|---|--|-------|
|                    |  |   | Mental Health Nursing, 23:537-551  |       |
| Shechtm<br>an 2010 | RQ 1.1-1.2 & 2.1-2.2 (searches combined) | Efficacy or safety data cannot be extracted | Shechtman Z, Mor M. Groups for children and adolescents with trauma-related symptoms: outcomes and processes. International journal of group psychotherapy. 2010 Apr;60(2):221-44. |       |

#### **Psychosocial: Psychoeducational interventions**

| Study ID       | Search             | Reason for exclusion               | Ref 1   | Ref 2 |
|----------------|--------------------|------------------------------------|---|-------|
| Ager<br>2011   | Handsearch         | Outcome measures are not validated | Ager A, Akesson B, Stark L, Flouri E, Okot B, McCollister F, Boothby N. The impact of the school-based Psychosocial Structured Activities (PSSA) program on conflict-affected children in northern Uganda. Journal of Child Psychology and Psychiatry. 2011 Nov 1;52(11):1124-33. |       |
| Kazdin<br>2002 | 2004 GL (excluded) | Commentary                         | Kazdin A.(2002) Comment on a school based psychosocial intervention was effective in children with persistent post-disaster trauma symptoms.) Evid Based Ment Health. 2002 Aug;5(3):76.   |       |

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Other non-pharmacological: Massage

| Study ID      | Search             | Reason for exclusion                        | Ref 1  | Ref 2 |
|---------------|--------------------|---|--|-------|
| Field<br>1996 | 2004 GL (excluded) | Efficacy or safety data cannot be extracted | Field, T., Seligman, S., Scafidi, F., & Schanberg, S. (1996). Alleviating posttraumatic stress in children following Hurricane Andrew. Journal of Applied Developmental Psychology, 17, Jan-Mar. |       |

#### **Economic studies**

No economic or utility studies were reviewed at full text and excluded from this review.

### **Appendix L – Research recommendations**

Research recommendations for "For children and young people with clinically important post-traumatic stress symptoms, what are the relative benefits and harms of psychological, psychosocial or other non-pharmacological interventions targeted at PTSD symptoms?"

No research recommendations were made for this review question.

#### Appendix M – Network Meta-Analysis: inconsistency checks

TSU, Bristol (Caitlin Daly and Sofia Dias)

#### Introduction

The purpose of this analysis was to critically assess the network meta-analysis (NMA) models used to estimate the comparative effectiveness of interventions for treating post-traumatic stress disorder (PTSD) in children and young people. The outcomes included in this analysis were 1) differences in PTSD symptoms recorded at baseline and end of treatment, 2) differences in PTSD symptoms recorded at baseline and 3-month follow-up, and 3) remission of PTSD.

#### Methods of network meta-analysis

#### **Conversion of Results Synthesized on Continuous Scale**

The economic model required probabilities of effect, which were informed by studies reporting continuous measures. To obtain these probabilities for the continuous outcomes, i.e. 1) differences in PTSD symptoms recorded at baseline and end of treatment and 2) differences in PTSD symptoms recorded at baseline and 3-month follow-up, the results of the evidence synthesis on the standardized mean difference (SMD) scale had to be transformed to a dichotomous scale. The log-odds ratio (LOR) of effect can be related to a notional SMD for effect using the formula (Chin, 2000; Higgins & Green, 2011):

$$LOR_{ck} = -\frac{\pi}{\sqrt{3}} SMD_{ck} \tag{1}$$

noting the change in sign to retain the interpretation of a positive LOR favouring treatment k.

The LORs were obtained by transforming the pooled treatment effects from the SMD scale using Equation (1).

#### **Model Critique**

When considering models for network meta-analysis (NMA), there are several aspects of the data that will impact the choice of parameters included in the model. Two important assumptions must be made in NMA regarding heterogeneity and consistency. Heterogeneity concerns the differences in treatment effects between trials within each treatment contrast, while consistency concerns the differences between the direct and indirect evidence informing the treatment contrasts (Dias 2011b & 2013b).

A fixed effect NMA model is the simplest model available to estimate the effects of interventions separately while simultaneously synthesizing all available evidence. This model assumes no heterogeneity between trials within each treatment contrast. In other words, all

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trials are estimating the same treatment effect, regardless of any differences in the conduct of the trials, populations, or treatments (i.e., administration or dose). If this assumption is unreasonable, then a random effects NMA model may be considered. This model accounts for any differences in treatment effects between trials that are beyond chance through measures such as the between-study standard deviation. When critiquing NMA models, it is good practice to assess and compare the fit of both fixed and random effects models, as differences may provide evidence of potential between-study heterogeneity.

Inconsistency was assessed by comparing the chosen consistency model (fixed or random effects) to an "inconsistency", or unrelated mean effects, model (Dias 2011b & 2013b). The latter is equivalent to having separate, unrelated, meta-analyses for every pairwise contrast, with a common variance parameter assumed in the case of random effects models. Note that inconsistency can only be assessed when there are closed loops of direct evidence on 3 treatments that are informed by at least 3 distinct trials (van Valkenhoef 2016).

The posterior mean of the residual deviance, which measures the magnitude of the differences between the observed data and the model predictions of the data, was used to assess the goodness of fit of each model (Spiegelhalter 2002). Smaller values are preferred, and in a well-fitting model the posterior mean residual deviance should be close to the number of data points in the network (each study arm contributes 1 data point) (Spiegelhalter 2002).

In addition to comparing how well the models fit the data using the posterior mean of the residual deviance, models were compared using the deviance information criterion (DIC). This is equal to the sum of the posterior mean deviance and the effective number of parameters, and thus penalizes model fit with model complexity (Spiegelhalter 2002). Lower values are preferred and typically differences of at least 3 points are considered meaningful (Spiegelhalter 2002).

The posterior mean between-study standard deviation, which measures the heterogeneity of treatment effects estimated by trials within contrasts, was also used to compare models. When comparing fixed and random effects models, it is important to assess whether there is enough evidence informing the between-study standard deviation. This was done by comparing the prior and posterior distributions of the between-study standard deviation. In addition, the magnitude of heterogeneity was considered. When comparing consistency and inconsistency models, if the inconsistency model has meaningfully smaller heterogeneity, then this indicates potential inconsistency in the data.

We performed further checks for evidence of inconsistency through node-splitting using the *gemtc* package in R (Dias 2010, 2011b & 2013b, van Valkenhoef 2016). This method permits the direct and indirect evidence contributing to an estimate of a relative effect to be split and compared (Dias 2010 & 2011b). To apply the node splitting method to the two continuous outcomes ('differences in PTSD symptoms recorded at baseline and end of treatment' and 'differences in PTSD symptoms recorded at baseline and 3-month follow-up') using the *gemtc* package, data were inputted at contrast level, where the SMDs of the treatment in arm *k* compared to the treatment in arm 1 for study *i* were calculated as

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$$SMD_{ik} = \frac{\bar{x}_{ik} - \bar{x}_{i1}}{SD_{pooled_i}}, \quad SD_{pooled_i} = \begin{cases} \sqrt{\frac{(n_{i1} - 1)sd_{i1}^2 + (n_{i2} - 1)sd_{i2}^2}{n_{i1} + n_{i2} - 2}} & \text{2-arm trial} \\ \sqrt{\frac{(n_{i1} - 1)sd_{i1}^2 + (n_{i2} - 1)sd_{i2}^2 + (n_{i3} - 1)sd_{i3}^2}{n_{i1} + n_{i2} + n_{i3} - 3}} & \text{3-arm trial} \end{cases}$$

with standard error

$$SE(SMD_{ik}) = \sqrt{Var(SMD_{ik})} \approx \begin{cases} \sqrt{\frac{1}{n_{i1}} + \frac{1}{n_{i2}} + \frac{SMD_{ik}^2}{2(n_{i1} + n_{i2} - 2)}} & \text{2-arm trial} \\ \sqrt{\frac{1}{n_{i1}} + \frac{1}{n_{ik}} + \frac{SMD_{ik}^2}{2(n_{i1} + n_{i2} + n_{i3} - 3)}} & \text{3-arm trial} \end{cases}$$

For trials with more than two arms, the *gemtc* package requires specification of the standard error of the mean of the baseline arm, as this determines the covariance of the differences. On a standardized scale, this is calculated as (Dias 2018):

$$se_{i1_{\text{standardized}}} = \frac{sd_{i1}}{SD_{pooled_i}\sqrt{n_{i1}}}$$
 .

#### Results

#### Outcome: Differences in PTSD symptoms recorded at baseline and end of treatment

Evidence on 17 interventions from 29 trials reporting the mean difference, standard deviation, and sample size across all intervention arms for this outcome was synthesized using NMA code provided by Dias for continuous data (Figure 188) (Dias 2018). Convergence was satisfactory for both fixed and random effects models (which assumed consistency) after 40,000 iterations, and the models were compared using results based on samples from a further 80,000 iterations on two chains. Large between trial heterogeneity was observed relative to the size of the intervention effect estimates ( $\tau$  (95% CrI) = 0.58 (0.37 – 0.89)) and the random effects model provided a better fit over the fixed effect model (Table 56).

Figure 188: Network diagram of comparisons for which direct evidence on differences in PTSD symptoms recorded at baseline and end of treatment was available.

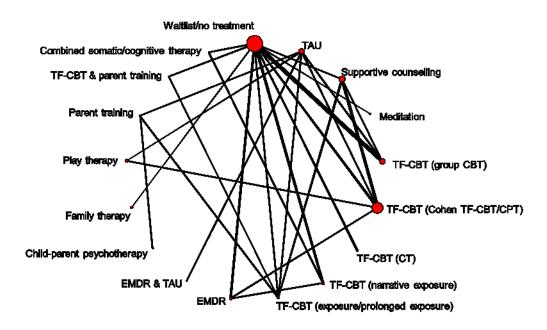


Table 56: Model fit statistics: differences in PTSD symptoms recorded at baseline and end of treatment

| Model  | Between Study Heterogeneity -<br>Standard Deviation (95% Crl) | Residual deviance <sup>a</sup> | DICb   |  |  |  |
|--|---|--------------------------------|--------|--|--|--|
| Fixed effect - consistency   | -   | 142.2                          | 340.17 |  |  |  |
| Random effects - consistency   | 0.58 (0.37 – 0.89)  | 63.01                          | 275.27 |  |  |  |
| Random effects - inconsistency   | 63.05   | 277.32                         |        |  |  |  |
| Posterior mean residual deviance compared to 63 total data points     Deviance information criteria (DIC) – lower values preferred |   |                                |        |  |  |  |

Since there were closed loops of direct evidence within the network that were informed by at least 3 distinct sets of trials, inconsistency checks were possible for this outcome. As the random effects model was preferred, a random effects inconsistency model was run. Convergence was satisfactory for this model after 40,000 iterations, and the consistency and inconsistency models were compared using results based on samples from a further 80,000 iterations on two chains. The WinBUGS code for the inconsistency model is provided in Appendix O.

No evidence of inconsistency was found through comparison of the consistency and inconsistency random effects models, as little difference was observed models (Table 56). The area below the line of equality in Figure 189 highlights where the inconsistency model better predicted data points, and the improvements were minimal. The additional parameters in the inconsistency model, which eliminates variation between treatment

contrasts, did not result in a decrease in the between-study heterogeneity (Table 56). Further checks for inconsistency using the node-splitting method (random effects model) did not find any evidence of inconsistency between the direct and indirect estimates (

Table 57, Figure 190).

Figure 189: Deviance contributions for the random effects consistency and inconsistency models: differences in PTSD symptoms recorded at baseline and end of treatment

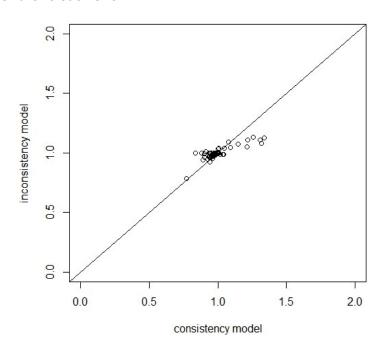


Table 57: Summary of node-splitting results: differences in PTSD symptoms recorded at baseline and end of treatment

| Node split model   | Heteroger | eity (SD)    | Residual | Data   | <b>p</b> -         |
|--|-----------|--------------|----------|--------|--------------------|
|  | Median    | 95% Crl      | deviance | points | value <sup>b</sup> |
| EMDR vs. Waitlist / no treatment   | 0.56      | (0.35, 0.93) | 33.36    | 33     | 0.63               |
| Supportive counselling vs. Waitlist / no treatment                               | 0.55      | (0.34, 0.91) | 33.6     | 33     | 0.61               |
| TF-CBT (group CBT) vs. Waitlist / no treatment                                   | 0.55      | (0.34, 0.92) | 33.5     | 33     | 0.62               |
| TF-CBT (Cohen TF-CBT / cognitive processing therapy) vs. Waitlist / no treatment | 0.53      | (0.33, 0.87) | 34.04    | 34     | 0.29               |
| TF-CBT (exposure/prolonged exposure) vs. Waitlist / no treatment                 | 0.54      | (0.33, 0.89) | 33.55    | 33     | 0.93               |
| TF-CBT (Cohen TF-CBT / cognitive processing therapy) vs. EMDR                    | 0.54      | (0.33, 0.89) | 34.38    | 34     | 0.64               |

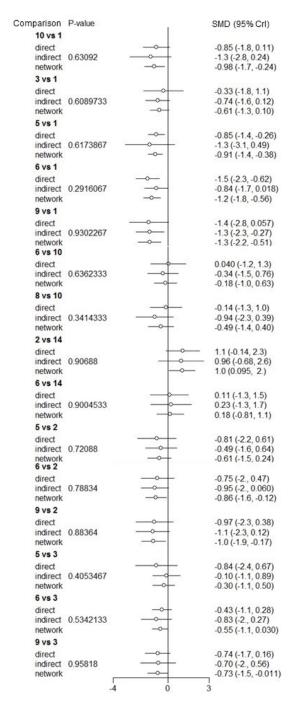
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| TF-CBT (narrative exposure) vs. EMDR  | 0.52 | (0.31, 0.87) | 33.3  | 33 | 0.34 |
|---|------|--------------|-------|----|------|
| TAU vs. Play therapy  | 0.55 | (0.34, 0.89) | 34.38 | 34 | 0.91 |
| TF-CBT (Cohen TF-CBT / cognitive processing therapy) vs. Play therapy           | 0.55 | (0.34, 0.89) | 34.39 | 34 | 0.90 |
| TF-CBT (group CBT) vs. TAU  | 0.54 | (0.33, 0.89) | 34.51 | 34 | 0.72 |
| TF-CBT (Cohen TF-CBT / cognitive processing therapy) vs. TAU                    | 0.55 | (0.34, 0.89) | 34.34 | 34 | 0.79 |
| TF-CBT (exposure/prolonged exposure) vs. TAU                                    | 0.54 | (0.33, 0.9)  | 33.42 | 33 | 0.88 |
| TF-CBT (group CBT) vs. Supportive counselling                                   | 0.55 | (0.34, 0.91) | 33.58 | 33 | 0.41 |
| TF-CBT (Cohen TF-CBT / cognitive processing therapy) vs. Supportive counselling | 0.54 | (0.33, 0.88) | 34.28 | 34 | 0.53 |
| TF-CBT (exposure/prolonged exposure) vs. Supportive counselling                 | 0.54 | (0.33, 0.89) | 34.48 | 34 | 0.96 |
| NMA (no nodes split)  | 0.52 | (0.32, 0.85) | 34.39 | 34 |      |

<sup>&</sup>lt;sup>a</sup> Number of data points to compare posterior mean residual deviance to

<sup>&</sup>lt;sup>b</sup> p-values < 0.05 is indicative of evidence of inconsistency between the direct and indirect estimates

Figure 190: Direct, indirect, and network estimates of relative treatment effects based on node-splitting results: differences in PTSD symptoms recorded at baseline and end of treatment



Treatments codes: 1 - Waitlist / no treatment, 2 - TAU, 3 - Supportive counselling, 4 - Meditation, 5 - TF-CBT (group CBT), 6 - TF-CBT (Cohen TF-CBT / cognitive processing therapy), 7 - TF-CBT (CT), 8 - TF-CBT (narrative exposure), 9 - TF-CBT (exposure/prolonged exposure), 10 - EMDR, 11 - EMDR

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& TAU, 12 - Child-parent psychotherapy, 13 - Family therapy, 14 - Play therapy, 15 - Parent training, 16 - TF-CBT & parent training, 17 - Combined somatic/cognitive therapy

In addition to the relative treatment effects estimated through NMA, we present direct and indirect estimates in the "Change Score\_Endpoint" worksheet in the "Supplementary File to Evidence Report B\_Appendix M" Excel file. The direct and indirect estimates are reported based on results given by the node-split models. All NMA estimates are reported based on the results from the random effects model that assumes consistency (Dias 2011a & 2013a).

#### Outcome: Differences in PTSD symptoms recorded at baseline and 3-month follow-up

Evidence on 12 interventions from 10 trials reporting the mean difference, standard deviation, and sample size across all intervention arms for this outcome was synthesised using NMA code for continuous data (Dias 2018). The network diagram is shown in Figure 191. Convergence was satisfactory for both fixed and random effects models (which assumed consistency) after 60,000 iterations, and the models were compared using results based on samples from a further 120,000 iterations on two chains. Large between trial heterogeneity was observed relative to the size of the intervention effect estimates ( $\tau$  (95% CrI) = 0.97 (0.30 – 2.69)). However, the distribution of the posterior between-study standard deviation suggests the prior distribution (Uniform(0,5)) has some influence on the estimate of heterogeneity (Figure 192). A sensitivity analysis with an informative prior should be conducted to assess whether this has an impact on the final results. Nevertheless, the random effects model provided a better fit over the fixed effect model (Table 58).

Figure 191: Network diagram of comparisons for which direct evidence on differences in PTSD symptoms recorded at baseline and 3-month follow-up was available.

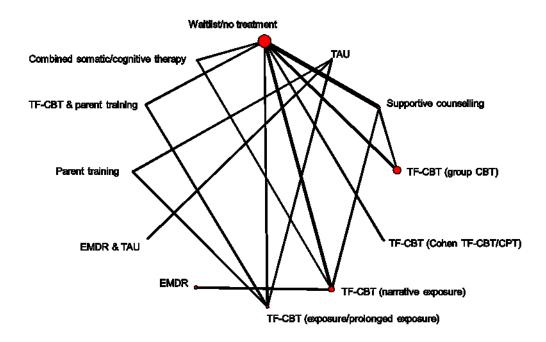


Figure 192: Density plot of the between-study standard deviation: differences in PTSD symptoms recorded at baseline and 3-month follow-up

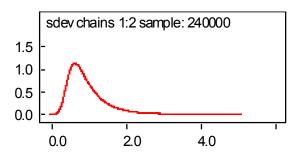


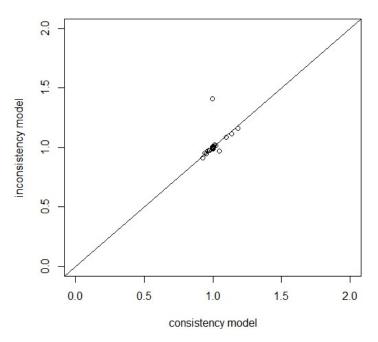
Table 58: Model fit statistics: differences in PTSD symptoms recorded at baseline and 3-month follow-up

| Model  | Between Study Heterogeneity  - Standard Deviation (95% Crl) | Residual deviance <sup>a</sup> | DICb   |  |
|--|---|--------------------------------|--------|--|
| Fixed effect – consistency   | -   | 41.51                          | 128.18 |  |
| Random effects – consistency   | 0.97 (0.30 – 2.69)  | 25.22                          | 115.57 |  |
| Random effects – inconsistency 0.90 (0.28 – 2.47) 25.11 115.39   |   |                                |        |  |
| <sup>a</sup> Posterior mean residual deviance compared to 25 total data points <sup>b</sup> Deviance information criteria (DIC) – lower values preferred |   |                                |        |  |

Since there were closed loops of direct evidence within the network that were informed by at least 3 distinct sets of trials, inconsistency checks were possible for this outcome. As the random effects model was preferred, a random effects inconsistency model was run. Convergence was satisfactory for this model after 60,000 iterations, and the consistency and inconsistency models were compared using results based on samples from a further 120,000 iterations on two chains. The WinBUGS code for the inconsistency model is provided in Appendix O.

There were no meaningful differences between posterior mean residual deviance of the consistency and inconsistency random effects models (Table 58, Figure 193). The inconsistency model provided poorer prediction for one data point compared to the consistency model (Figure 193). The inconsistency model has slightly smaller between-study heterogeneity than the consistency model, although this was still large (Table 58). However, as previously noted, the between-study heterogeneity is poorly estimated from the data, and results should be interpreted with caution.

Figure 193: Deviance contributions for the random effects consistency and inconsistency models: differences in PTSD symptoms recorded at baseline and 3-month follow-up



Further checks for inconsistency using the node-splitting method (random effects model) did not find any evidence of inconsistency between the direct and indirect estimates (Table 59, Figure 194). In addition to the relative treatment effects estimated through NMA, we present direct and indirect estimates in the "Supplementary File to Evidence Report B\_Appendix M" Excel file. The direct and indirect estimates are reported based on results given by the node-split models. All NMA estimates are reported based on the results from the random effects model that assumes consistency (Dias 2011a & 2013a). There is considerable uncertainty in the results of treatment effect estimates, and this is due to the small size of the included

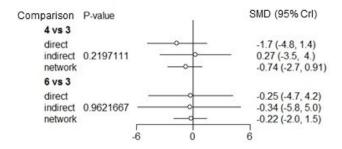
studies, the small total number of studies, and the large heterogeneity that is present. Thus, caution should be exercised when interpreting the results.

Table 59: Summary of node-splitting results: differences in PTSD symptoms recorded at baseline and 3-month follow-up

| Node split model  | Heterogeneity (SD) |              | Residual | Data                | p-value <sup>b</sup> |
|---|--------------------|--------------|----------|---------------------|----------------------|
|   | median             | 95% CrI      | deviance | points <sup>a</sup> |                      |
| TF-CBT (group CBT) vs. Supportive counselling                                     | 0.72               | (0.09, 3.92) | 14.17    | 14                  | 0.22                 |
| TF-CBT (narrative exposure) vs. Supportive counselling                            | 1.44               | (0.35, 4.50) | 14.21    | 14                  | 0.96                 |
| NMA (no nodes split)  | 0.74               | (0.22, 2.62) | 15.46    | 15                  |                      |
| <sup>a</sup> Number of data points to compare posterior mean residual deviance to |                    |              |          |                     |                      |

<sup>b</sup>p-values < 0.05 is indicative of evidence of inconsistency between the direct and indirect estimates

Figure 194: Direct, indirect, and network estimates of relative treatment effects based on node-splitting results: differences in PTSD symptoms recorded at baseline and 3-month follow-up



Treatment codes: 1 - Waitlist / no treatment, 2 - TAU, 3 - Supportive counselling, 4 - TF-CBT (group CBT), 5 - TF-CBT (Cohen TF-CBT / cognitive processing therapy), 6 - TF-CBT (narrative exposure), 7 - TF-CBT (exposure/prolonged exposure), 8 - EMDR, 9 - EMDR & TAU, 10 - Parent training, 11 - TF-CBT & parent training, 12 - Combined somatic/cognitive therapy

#### **Outcome: Remission**

Evidence on 7 interventions from 9 trials reporting the number of participants who achieved remission from PTSD and sample size across all intervention arms for this outcome was synthesized using the NMA code provided by Dias and colleagues for binary data (Dias 2011a & 2013a). The network of evidence is shown in Figure 195. Convergence was satisfactory for both fixed and random effects models (which assumed consistency) after 40,000 iterations, and the models were compared using results based on samples from a further 80,000 iterations on two chains.

Large between trial heterogeneity was observed relative to the size of the intervention effect estimates ( $\tau$  (95% CrI) = 0.65 (0.03 – 1.80)). However, the distribution of the posterior

between-study standard deviation suggests the prior distribution (Uniform(0,2)) has a heavy influence on the estimate of heterogeneity (Figure 196). This is because there is very little evidence to inform the between-study heterogeneity. A sensitivity analysis with an informative prior should be conducted to assess whether this has an impact on the final results. Nevertheless, the fixed and random effects models both provided good fit and no meaningful differences were observed between the fit of both models (Table 60). Therefore, the simpler, fixed effect model was chosen.

Figure 195: Network diagram of comparisons for which direct evidence on remission from PTSD was available: remission

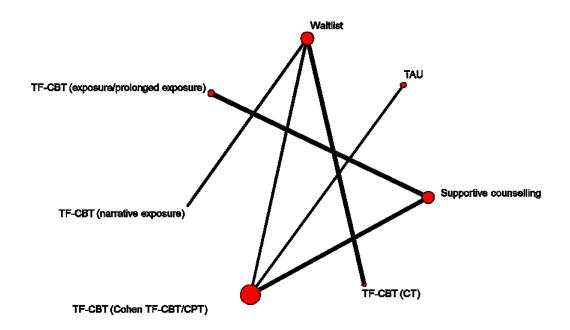


Figure 196: Density plot of the between-study standard deviation: remission

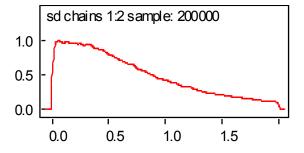


Table 60: Model fit statistics: remission

| Model  | Between Study Heterogeneity -<br>Standard Deviation (95% Crl) | Residual deviance <sup>a</sup> | DICb  |  |  |
|--|---|--------------------------------|-------|--|--|
| Fixed effect - consistency   | -   | 17.37                          | 93.71 |  |  |
| Random effects - consistency 0.65 (0.03 – 1.80) 17.38 95.0   |   |                                |       |  |  |
| a Posterior mean residual deviance compared to 18 total data points b Deviance information criteria (DIC) – lower values preferred |   |                                |       |  |  |

Since there were no closed loops of direct evidence within the network (Figure 195), inconsistency checks were not possible for this outcome.

#### Conclusion

The inconsistency checks did not identify any evidence of inconsistency in the direct and indirect evidence included in the network meta-analyses. However, the small amount of evidence informing the 'differences in PTSD symptoms recorded at baseline and 3-month follow-up' and 'remission of PTSD' outcomes is reflected by the large uncertainty of the results. For example, the odds ratio and 95% credible interval of TF-CBT (narrative exposure) compared to waitlist in the remission outcome is 15.58 (2.44 – 163.1). The large amount of uncertainty in these results decreases the reliability of the probability of a treatment ranking best and thus it is not a suitable measure to base decisions on (Kibret 2014). Caution must be exercised when interpreting the results.

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# Appendix N – additional results of Network Meta-Analysis described in Appendix J (all pair-wise comparisons)

## PTSD symptom scores, change from baseline to endpoint: Standardised Mean Differences (SMD)

[negative values favour second intervention in the comparison]

| Comparison | Effect (mean SMD, 95% Crl) | Comparison | Effect (mean SMD, 95% Crl) |
|------------|----------------------------|------------|----------------------------|
| diff[2,3]  | -0.289 (-1.175 to 0.593)   | diff[3,4]  | -1.074 (-2.533 to 0.343)   |
| diff[2,4]  | -1.363 (-2.883 to 0.162)   | diff[3,5]  | -0.314 (-1.127 to 0.502)   |
| diff[2,5]  | -0.603 (-1.509 to 0.311)   | diff[3,6]  | -0.574 (-1.179 to 0.0249)  |
| diff[2,6]  | -0.864 (-1.653 to -0.074)  | diff[3,7]  | -2.343 (-3.562 to -1.155)  |
| diff[2,7]  | -2.632 (-3.973 to -1.329)  | diff[3,8]  | -0.899 (-1.924 to 0.110)   |
| diff[2,8]  | -1.188 (-2.320 to -0.072)  | diff[3,9]  | -0.742 (-1.490 to 0.004)   |
| diff[2,9]  | -1.031 (-1.941 to -0.124)  | diff[3,10] | -0.395 (-1.366 to 0.545)   |
| diff[2,10] | -0.685 (-1.782 to 0.391)   | diff[3,11] | 0.310 (-1.398 to 2.027)    |
| diff[2,11] | 0.021 (-1.445 to 1.479)    | diff[3,12] | -1.567 (-3.438 to 0.295)   |
| diff[2,12] | -1.856 (-3.634 to -0.079)  | diff[3,13] | 0.225 (-1.184 to 1.619)    |
| diff[2,13] | -0.067 (-1.591 to 1.407)   | diff[3,14] | -0.750 (-1.914 to 0.399)   |
| diff[2,14] | -1.039 (-2.043 to -0.061)  | diff[3,15] | -0.366 (-1.710 to 0.996)   |
| diff[2,15] | -0.655 (-1.923 to 0.595)   | diff[3,16] | -1.194 (-2.604 to 0.249)   |
| diff[2,16] | -1.483 (-3.011 to 0.025)   | diff[3,17] | -1.542 (-2.934 to -0.150)  |
| diff[2,17] | -1.831 (-3.295 to -0.361)  |            |                            |
|            |                            |            |                            |
| diff[4,5]  | 0.760 (-0.620 to 2.147)    | diff[5,6]  | -0.260 (-1.030 to 0.514)   |
| diff[4,6]  | 0.500 (-0.887 to 1.905)    | diff[5,7]  | -2.029 (-3.182 to -0.885)  |
| diff[4,7]  | -1.269 (-2.874 to 0.335)   | diff[5,8]  | -0.585 (-1.524 to 0.355)   |
| diff[4,8]  | 0.175 (-1.286 to 1.637)    | diff[5,9]  | -0.428 (-1.363 to 0.503)   |
| diff[4,9]  | 0.332 (-1.167 to 1.834)    | diff[5,10] | -0.081 (-1.020 to 0.856)   |
| diff[4,10] | 0.679 (-0.771 to 2.166)    | diff[5,11] | 0.624 (-1.079 to 2.340)    |
| diff[4,11] | 1.384 (-0.743 to 3.525)    | diff[5,12] | -1.253 (-3.146 to 0.663)   |
| diff[4,12] | -0.493 (-2.756 to 1.781)   | diff[5,13] | 0.537 (-0.829 to 1.869)    |
| diff[4,13] | 1.296 (-0.468 to 3.015)    | diff[5,14] | -0.436 (-1.617 to 0.777)   |
| diff[4,14] | 0.324 (-1.355 to 2.003)    | diff[5,15] | -0.0515 (-1.470 to 1.369)  |
| diff[4,15] | 0.708 (-1.130 to 2.585)    | diff[5,16] | -0.880 (-2.329 to 0.566)   |
| diff[4,16] | -0.120 (-1.969 to 1.712)   | diff[5,17] | -1.228 (-2.547 to 0.114)   |
| diff[4,17] | -0.468 (-2.219 to 1.279)   |            |                            |
|            |                            |            |                            |
| diff[6,7]  | -1.768 (-2.949 to -0.615)  | diff[7,8]  | 1.444 (0.183 to 2.688)     |
| diff[6,8]  | -0.325 (-1.276 to 0.616)   | diff[7,9]  | 1.601 (0.331 to 2.900)     |

|             | ·                                     |             |                           |
|-------------|---------------------------------------|-------------|---------------------------|
| diff[6,9]   | -0.168 (-0.986 to 0.642)              | diff[7,10]  | 1.947 (0.687 to 3.232)    |
| diff[6,10]  | 0.179 (-0.674 to 1.031)               | diff[7,11]  | 2.652 (0.697 to 4.623)    |
| diff[6,11]  | 0.884 (-0.759 to 2.525)               | diff[7,12]  | 0.776 (-1.353 to 2.893)   |
| diff[6,12]  | -0.993 (-2.85 to 0.854)               | diff[7,13]  | 2.565 (0.972 to 4.139)    |
| diff[6,13]  | 0.797 (-0.578 to 2.159)               | diff[7,14]  | 1.593 (0.074 to 3.120)    |
| diff[6,14]  | -0.176 (-1.205 to 0.868)              | diff[7,15]  | 1.977 (0.280 to 3.691)    |
| diff[6,15]  | 0.209 (-1.134 to 1.559)               | diff[7,16]  | 1.149 (-0.545 to 2.828)   |
| diff[6,16]  | -0.620 (-2.043 to 0.798)              | diff[7,17]  | 0.8010 (-0.746 to 2.385)  |
| diff[6,17]  | -0.967 (-2.305 to 0.374)              |             |                           |
|             |                                       |             |                           |
| diff[8,9]   | 0.157 (-0.938 to 1.251)               | diff[9,10]  | 0.347 (-0.710 to 1.417)   |
| diff[8,10]  | 0.504 (-0.425 to 1.447)               | diff[9,11]  | 1.052 (-0.661 to 2.793)   |
| diff[8,11]  | 1.209 (-0.628 to 3.049)               | diff[9,12]  | -0.825 (-2.608 to 0.968)  |
| diff[8,12]  | -0.668 (-2.671 to 1.356)              | diff[9,13]  | 0.964 (-0.523 to 2.437)   |
| diff[8,13]  | 1.121 (-0.329 to 2.563)               | diff[9,14]  | -0.008 (-1.219 to 1.220)  |
| diff[8,14]  | 0.149 (-1.205 to 1.488)               | diff[9,15]  | 0.376 (-0.857 to 1.617)   |
| diff[8,15]  | 0.533 (-1.005 to 2.103)               | diff[9,16]  | -0.452 (-1.83 to 0.919)   |
| diff[8,16]  | -0.295 (-1.811 to 1.247)              | diff[9,17]  | -0.800 (-2.254 to 0.639)  |
| diff[8,17]  | -0.643 (-1.831 to 0.566)              |             |                           |
|             |                                       |             |                           |
| diff[10,11] | 0.705 (-1.104 to 2.520)               | diff[11,12] | -1.877 (-4.180 to 0.426)  |
| diff[10,12] | -1.172 (-3.151 to 0.825)              | diff[11,13] | -0.087 (-2.200 to 2.007)  |
| diff[10,13] | 0.618 (-0.817 to 2.052)               | diff[11,14] | -1.060 (-2.826 to 0.719)  |
| diff[10,14] | -0.355 (-1.668 to 0.927)              | diff[11,15] | -0.675 (-2.635 to 1.258)  |
| diff[10,15] | 0.030 (-1.500 to 1.550)               | diff[11,16] | -1.504 (-3.609 to 0.624)  |
| diff[10,16] | -0.799 (-2.342 to 0.728)              | diff[11,17] | -1.852 (-3.941 to 0.249)  |
| diff[10,17] | -1.146 (-2.532 to 0.221)              |             |                           |
|             |                                       |             |                           |
| diff[12,13] | 1.789 (-0.462 to 4.045)               | diff[13,14] | -0.973 (-2.640 to 0.702)  |
| diff[12,14] | 0.817 (-1.170 to 2.812)               | diff[13,15] | -0.588 (-2.432 to 1.258)  |
| diff[12,15] | 1.201 (-0.088 to 2.475)               | diff[13,16] | -1.417 (-3.223 to 0.412)  |
| diff[12,16] | 0.373 (-1.848 to 2.551)               | diff[13,17] | -1.764 (-3.479 to -0.040) |
| diff[12,17] | 0.025 (-2.211 to 2.211)               |             |                           |
|             |                                       |             |                           |
| diff[14,15] | 0.385 (-1.171 to 1.905)               | diff[15,16] | -0.829 (-2.611 to 0.980)  |
| diff[14,16] | -0.444 (-2.156 to 1.270)              | diff[15,17] | -1.176 (-3.002 to 0.618)  |
| diff[14,17] | -0.792 (-2.437 to 0.842)              |             |                           |
|             |                                       |             |                           |
| diff[16,17] | -0.3476 (-2.18 to 1.458)              |             |                           |
|             | · · · · · · · · · · · · · · · · · · · |             |                           |

# PTSD symptom scores, change from baseline to 1-4 months follow-up: Standardised Mean Differences (SMD)

[negative values favour second intervention in the comparison]

| Comparison  | Effect (mean SMD, 95% Crl) | Comparison  | Effect (mean SMD, 95% Crl) |
|-------------|----------------------------|-------------|----------------------------|
| diff[2,3]   | -0.390 (-2.351 to 1.740)   | diff[3,4]   | -0.776 (-1.930 to 0.244)   |
| diff[2,4]   | -1.166 (-3.315 to 0.923)   | diff[3,5]   | -1.006 (-2.590 to 0.446)   |
| diff[2,5]   | -1.396 (-3.695 to 0.904)   | diff[3,6]   | -0.205 (-1.279 to 0.742)   |
| diff[2,6]   | -0.595 (-2.654 to 1.529)   | diff[3,7]   | -0.186 (-1.759 to 1.253)   |
| diff[2,7]   | -0.576 (-1.958 to 0.789)   | diff[3,8]   | 0.144 (-1.521 to 1.698)    |
| diff[2,8]   | -0.247 (-2.642 to 2.209)   | diff[3,9]   | -0.362 (-2.944 to 2.050)   |
| diff[2,9]   | -0.752 (-2.193 to 0.683)   | diff[3,10]  | -0.303 (-2.383 to 1.603)   |
| diff[2,10]  | -0.693 (-2.097 to 0.690)   | diff[3,11]  | -0.750 (-2.411 to 0.778)   |
| diff[2,11]  | -1.141 (-3.170 to 0.901)   | diff[3,12]  | -1.061 (-2.503 to 0.238)   |
| diff[2,12]  | -1.452 (-3.651 to 0.810)   |             |                            |
|             |                            |             |                            |
| diff[4,5]   | -0.230 (-1.861 to 1.412)   | diff[5,6]   | 0.801 (-0.8054 to 2.394)   |
| diff[4,6]   | 0.571 (-0.660 to 1.885)    | diff[5,7]   | 0.820 (-1.003 to 2.692)    |
| diff[4,7]   | 0.590 (-1.026 to 2.233)    | diff[5,8]   | 1.150 (-0.847 to 3.201)    |
| diff[4,8]   | 0.920 (-0.8327 to 2.72)    | diff[5,9]   | 0.644 (-2.057 to 3.327)    |
| diff[4,9]   | 0.415 (-2.123 to 2.972)    | diff[5,10]  | 0.703 (-1.558 to 2.943)    |
| diff[4,10]  | 0.473 (-1.596 to 2.554)    | diff[5,11]  | 0.256 (-1.687 to 2.187)    |
| diff[4,11]  | 0.026 (-1.658 to 1.720)    | diff[5,12]  | -0.055 (-1.872 to 1.779)   |
| diff[4,12]  | -0.285 (-1.752 to 1.281)   |             |                            |
|             |                            |             |                            |
| diff[6,7]   | 0.019 (-1.570 to 1.579)    | diff[7,8]   | 0.330 (-1.695 to 2.381)    |
| diff[6,8]   | 0.348 (-0.920 to 1.612)    | diff[7,9]   | -0.176 (-2.151 to 1.828)   |
| diff[6,9]   | -0.157 (-2.72 to 2.338)    | diff[7,10]  | -0.117 (-1.454 to 1.172)   |
| diff[6,10]  | -0.098 (-2.169 to 1.888)   | diff[7,11]  | -0.564 (-2.022 to 0.903)   |
| diff[6,11]  | -0.546 (-2.234 to 1.128)   | diff[7,12]  | -0.875 (-2.658 to 0.900)   |
| diff[6,12]  | -0.857 (-2.069 to 0.366)   |             |                            |
|             |                            |             |                            |
| diff[8,9]   | -0.505 (-3.323 to 2.273)   | diff[9,10]  | 0.059 (-1.974 to 2.058)    |
| diff[8,10]  | -0.447 (-2.854 to 1.918)   | diff[9,11]  | -0.389 (-2.873 to 2.112)   |
| diff[8,11]  | -0.894 (-2.974 to 1.173)   | diff[9,12]  | -0.700 (-3.330 to 1.988)   |
| diff[8,12]  | -1.205 (-2.954 to 0.564)   |             |                            |
|             |                            |             |                            |
| diff[10,11] | -0.447 (-2.386 to 1.551)   | diff[11,12] | -0.311 (-2.164 to 1.554)   |
| diff[10,12] | -0.758 (-2.948 to 1.433)   |             |                            |

# DRAFT FOR CONSULTATION Psychological, psychosocial and other non-pharmacological interventions for the treatment of PTSD in children and young people

#### Remission (loss of PTSD diagnosis according to ICD/DCM criteria or similar): logodds ratios

[positive values favour second intervention in the comparison]

| Comparison | Effect (mean, 95% Crl) | Comparison | Effect (mean, 95% Crl) |
|------------|------------------------|------------|------------------------|
| lor[2,3]   | 0.36 (-0.95 to 1.69)   | lor[3,4]   | 2.51 (0.72 to 4.44)    |
| lor[2,4]   | 2.87 (1.01 to 4.88)    | lor[3,5]   | 0.74 (-0.11 to 1.60)   |
| lor[2,5]   | 1.10 (0.13 to 2.15)    | lor[3,6]   | 2.66 (0.40 to 5.18)    |
| lor[2,6]   | 3.02 (0.71 to 5.62)    | lor[3,7]   | 1.47 (0.62 to 2.36)    |
| lor[2,7]   | 1.83 (0.26 to 3.45)    |            |                        |
|            |                        |            |                        |
| lor[4,5]   | -1.77 (-3.49 to -0.20) | lor[5,6]   | 1.92 (-0.16 to 4.31)   |
| lor[4,6]   | 0.15 (-2.33 to 2.83)   | lor[5,7]   | 0.73 (-0.49 to 1.96)   |
| lor[4,7]   | -1.04 (-3.14 to 0.95)  |            |                        |
|            | •                      |            |                        |
| lor[6,7]   | -1.19 (-3.83 to 1.27)  |            |                        |

# Appendix O – WinBUGS code for inconsistency model described in Appendix M – 'Changes in PTSD Symptom Scores between Baseline and Treatment Endpoint' and 'Changes in PTSD Symptom Scores between Baseline and 1-4 Month Follow-Up'

```
# Normal likelihood, identity link: SMD with arm-based means;
# Random effects inconsistency model
                           # *** PROGRAM STARTS
model{
for(i in 1:ns){
                            # LOOP THROUGH STUDIES
 delta[i,1] <- 0
                           # treatment effect is zero for control arm
 mu[i] \sim dnorm(0,.0001)
                                  # vague priors for all trial baselines
}
# CONTINUOUS DATA AS ARM MEANS
for(i in 1:ns){
 # calculate pooled.sd and adjustment for SMD
 df[i] <- sum(n[i,1:na[i]]) - na[i] # denominator for pooled.var
 Pooled.var[i] <- sum(nvar[i,1:na[i]])/df[i]
 Pooled.sd[i] <- sqrt(Pooled.var[i]) # pooled sd for study i, for SMD
# H[i] <- 1 - 3/(4*df[i]-1)
                               # use Hedges' g
 H[i] <- 1
                          # use Cohen's d (ie no adjustment)
 for (k in 1:na[i]){
  se[i,k] \leftarrow sd[i,k]/sqrt(n[i,k])
  var[i,k] \leftarrow pow(se[i,k],2)
                               # calcultate variances
  prec[i,k] <- 1/var[i,k]
                             # set precisions
  y[i,k] ~ dnorm(phi[i,k], prec[i,k]) # normal likelihood
  phi[i,k] <- theta[i,k] * (Pooled.sd[i]/H[i]) # theta is standardised mean
  theta[i,k] <- mu[i] + delta[i,k] # model for linear predictor, delta is SMD
  dev[i,k] \leftarrow (y[i,k]-phi[i,k])*(y[i,k]-phi[i,k])*prec[i,k]
  nvar[i,k] \leftarrow (n[i,k]-1) * pow(sd[i,k],2) # for pooled.sd
```

```
}
 # summed residual deviance contribution for this trial
 resdev[i] <- sum(dev[i,1:na[i]])
}
# RE MODEL
for(i in 1:ns){
                           #LOOP THROUGH ALL STUDIES
 for (k in 2:na[i]){
                           # LOOP THROUGH ARMS
  # trial-specific RE distributions
  delta[i,k] \sim dnorm(d[t[i,1], t[i,k]], tau)
  }
}
#
totresdev <- sum(resdev[]) # Total Residual Deviance (all data)
# Priors distributions
sdev \sim dunif(0,5)
                   # vague prior for between-trial SD
tau <- pow(sdev,-2)
                               # between-trial precision
# vague prior for treatment effects
for (c in 1:(nt-1)){
 d[c,c]<-0
 for (k \text{ in } (c+1):nt) \{ d[c,k] \sim dnorm(0,.001) \}
}
}
                     # *** PROGRAM ENDS
```