

Author(s): NCCMH

Date: 2008-12-02

Question: Should drugs (versus placebo) (efficacy data) be used for subthreshold depression?

Settings:

Bibliography:

Quality assessment							Summary of findings				Quality	Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients drugs (versus placebo) (efficacy data)	control	Relative (95% CI)	Absolute		
Number of people not achieving at least 50% reduction in depression score - SSRIs: dysthymia only												
5	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	176/382 (46.1%)	223/345 (64.6%)	RR 0.72 (0.63 to 0.82)	18 fewer per 100 (from 12 fewer to 24 fewer)	⊕⊕⊕⊕ HIGH	
								66.5%		19 fewer per 100 (from 12 fewer to 25 fewer)		
Number of people not achieving at least 50% reduction in depression score - SSRIs: minor depression only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	55/106 (51.9%)	57/109 (52.3%)	RR 0.99 (0.77 to 1.28)	1 fewer per 100 (from 12 fewer to 15 more)	⊕⊕⊕⊕ MODERATE	
								52.3%		1 fewer per 100 (from 12 fewer to 15 more)		
Number of people not achieving at least 50% reduction in depression score - TCAs: dysthymia only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	25/68 (36.8%)	54/76 (71.1%)	RR 0.52 (0.37 to 0.73)	34 fewer per 100 (from 19 fewer to 45 fewer)	⊕⊕⊕⊕ MODERATE	
								71.1%		34 fewer per 100 (from 19 fewer to 45 fewer)		
Number of people not achieving at least 50% reduction in depression score - MAIOs: dysthymia only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ³	none	25/70 (35.7%)	54/76 (71.1%)	RR 0.5 (0.36 to 0.71)	36 fewer per 100 (from 21 fewer to 45 fewer)	⊕⊕⊕⊕ MODERATE	
								71.1%		36 fewer per 100 (from 21 fewer to 46 fewer)		
Number of people not achieving remission - SSRIs: dysthymia only												
3	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	167/317 (52.7%)	194/291 (66.7%)	RR 0.78 (0.68 to 0.89)	15 fewer per 100 (from 7 fewer to 21 fewer)	⊕⊕⊕⊕ HIGH	
								67.9%		15 fewer per 100 (from 7 fewer to 22 fewer)		
Number of people not achieving remission - SSRIs: minor depression only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁴	none	62/106 (58.5%)	60/109 (55%)	RR 1.06 (0.84 to 1.34)	3 more per 100 (from 9 fewer to 19 more)	⊕⊕⊕⊕ MODERATE	
								55.1%		3 more per 100 (from 9 fewer to 19 more)		
Number of people not achieving remission - TCAs: dysthymia only												
2	randomised trials	no serious limitations	serious ⁵	no serious indirectness	serious ⁶	none	120/204 (58.8%)	154/216 (71.3%)	RR 0.81 (0.63 to 1.03)	14 fewer per 100 (from 26 fewer to 2 more)	⊕⊕⊕⊕ LOW	
								72.7%		14 fewer per 100 (from 27 fewer to 2 more)		
Number of people not achieving remission - MAIOs: dysthymia only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁷	none	34/70 (48.6%)	59/76 (77.6%)	RR 0.63 (0.48 to 0.82)	29 fewer per 100 (from 14 fewer to 40 fewer)	⊕⊕⊕⊕ MODERATE	
								77.6%		29 fewer per 100 (from 14 fewer to 40 fewer)		
Mean endpoint scores (clinician rated) - SSRIs: dysthymia only (Better indicated by lower values)												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	104	115	-	SMD 0.56 lower (0.83 to 0.29 lower)	⊕⊕⊕⊕ HIGH	
Mean endpoint scores (clinician rated) - SSRIs: minor depression only (Better indicated by lower values)												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁸	none	155	167	-	SMD 0.19 lower (0.41 lower to 0.03 higher)	⊕⊕⊕⊕ MODERATE	
Mean endpoint scores (clinician rated) - TCAs: dysthymia only (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁹	none	107	105	-	SMD 0.62 lower (0.9 to 0.35 lower)	⊕⊕⊕⊕ MODERATE	

Mean endpoint scores (clinician rated) - APs: dysthymia only (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁰	none	101	105	-	SMD 0.66 lower (0.94 to 0.38 lower)	⊕⊕⊕⊕ MODERATE	
Mean endpoint scores (self rated) (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹¹	none	73	74	-	SMD 0.4 lower (0.72 to 0.07 lower)	⊕⊕⊕⊕ MODERATE	
Mean endpoint scores (self rated) - SSRIs: minor depression only (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹²	none	73	74	-	SMD 0.4 lower (0.72 to 0.07 lower)	⊕⊕⊕⊕ MODERATE	
Mean change (clinician rated) - SSRIs: dysthymia only (Better indicated by lower values)												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	206	179	-	SMD 0.31 lower (0.51 to 0.11 lower)	⊕⊕⊕⊕ HIGH	
Mean change (clinician rated) - TCAs: dysthymia only (Better indicated by lower values)												
3	randomised trials	no serious limitations	serious ¹³	no serious indirectness	no serious imprecision	none	306	317	-	SMD 0.61 lower (0.9 to 0.31 lower)	⊕⊕⊕⊕ MODERATE	
Mean change (clinician rated) - APs: dysthymia only (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁴	none	101	105	-	SMD 0.67 lower (0.95 to 0.39 lower)	⊕⊕⊕⊕ MODERATE	
Mean change (clinician rated) - MAOIs: dysthymia only (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁵	none	67	72	-	SMD 0.97 lower (1.32 to 0.62 lower)	⊕⊕⊕⊕ MODERATE	

- ¹ Single study
- ² As 1
- ³ As 1
- ⁴ As 1
- ⁵ Significant heterogeneity - random effects model used
- ⁶ Non significant effect size
- ⁷ As 1
- ⁸ Non significant effect size
- ⁹ As 1
- ¹⁰ As 1
- ¹¹ As 1
- ¹² As 1
- ¹³ As 5
- ¹⁴ As 1
- ¹⁵ As 1

Author(s): NCCMH
Date: 2008-12-02
Question: Should drugs (versus placebo) (acceptability/tolerability data) be used for subthreshold depression?
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Quality assessment							Summary of findings				Quality	Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect			
							drugs (versus placebo) (acceptability/tolerability data)	control	Relative (95% CI)	Absolute		
Leaving the study early - SSRIs: dysthymia only												
6	randomised trials	no serious limitations	serious ¹	no serious indirectness	serious ²	none	101/535 (18.9%)	108/495 (21.8%)	RR 0.84 (0.57 to 1.24)	3 fewer per 100 (from 9 fewer to 5 more)	⊕⊕⊕⊕ LOW	
								21.5%		3 fewer per 100 (from 9 fewer to 5 more)		
Leaving the study early - SSRIs: minor depression												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ³	none	59/187 (31.6%)	50/190 (26.3%)	RR 1.2 (0.87 to 1.65)	5 more per 100 (from 3 fewer to 17 more)	⊕⊕⊕⊕ LOW	
								26.4%		5 more per 100 (from 3 fewer to 17 more)		
Leaving the study early - TCAs: dysthymia only												
4	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁴	none	85/366 (23.2%)	78/368 (21.2%)	RR 1.1 (0.84 to 1.44)	2 more per 100 (from 3 fewer to 9 more)	⊕⊕⊕⊕ MODERATE	
								22.3%		2 more per 100 (from 4 fewer to 10 more)		
Leaving the study early - MAOIs: dysthymia only												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁵	none	13/108 (12%)	15/104 (14.4%)	RR 0.83 (0.42 to 1.24)	2 fewer per 100 (from 8 fewer to 10 more)	⊕⊕⊕⊕ LOW	
										2 fewer		

									14.4%	1.67	per 100 (from 8 fewer to 10 more)		
Leaving the study early - APs: dysthymia only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁶	none		14/104 (13.5%)	22/108 (20.4%) 20.4%	RR 0.66 (0.36 to 1.22)	7 fewer per 100 (from 13 fewer to 4 more) 7 fewer per 100 (from 13 fewer to 4 more)	⊕⊕⊕○	MODERATE
Leaving the study early due to side effects - SSRIs: dysthymia only													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁷	none		12/245 (4.9%)	7/252 (2.8%) 2.7%	RR 1.77 (0.71 to 4.41)	2 more per 100 (from 1 fewer to 9 more) 2 more per 100 (from 1 fewer to 9 more)	⊕⊕⊕○	MODERATE
Leaving the study early due to side effects - SSRIs: minor depression only													
2	randomised trials	no serious limitations	serious ⁸	no serious indirectness	very serious ⁹	none		17/187 (9.1%)	10/190 (5.3%) 5.2%	RR 1.55 (0.51 to 4.68)	3 more per 100 (from 3 fewer to 19 more) 3 more per 100 (from 3 fewer to 19 more)	⊕○○○	VERY LOW
Leaving the study early due to side effects - TCAs: dysthymia only													
4	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none		45/366 (12.3%)	8/369 (2.2%) 1.4%	RR 5.44 (2.66 to 11.11)	10 more per 100 (from 4 more to 22 more) 6 more per 100 (from 2 more to 14 more)	⊕⊕⊕⊕	HIGH
Leaving the study early due to side effects - MAOIs: dysthymia only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁰	none		7/108 (6.5%)	2/104 (1.9%) 1.9%	RR 3.37 (0.72 to 15.85)	5 more per 100 (from 1 fewer to 29 more) 5 more per 100 (from 1 fewer to 28 more)	⊕⊕⊕○	MODERATE
Leaving the study early due to side effects - APs: dysthymia only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹¹	none		3/104 (2.9%)	1/108 (0.9%) 0.9%	RR 3.12 (0.33 to 29.47)	2 more per 100 (from 1 fewer to 26 more) 2 more per 100 (from 1 fewer to 26 more)	⊕⊕○○	LOW
Patients reporting side effects - SSRIs: dysthymia only													
3	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none		188/360 (52.2%)	153/313 (48.9%) 44.9%	RR 1.09 (0.95 to 1.25)	4 more per 100 (from 2 fewer to 12 more) 4 more per 100 (from 2 fewer to 11 more)	⊕⊕⊕⊕	HIGH
Patients reporting side effects - SSRIs: minor depression only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹²	none		25/106 (23.6%)	34/109 (31.2%) 31.2%	RR 0.76 (0.49 to 1.18)	7 fewer per 100 (from 16 fewer to 6 more) 7 fewer per 100 (from 16 fewer to 6 more)	⊕⊕⊕○	MODERATE
Patients reporting side effects - TCAs: dysthymia only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹³	none		69/111 (62.2%)	48/108 (44.4%) 18 more	RR 1.4 (1.08 to 1.81)	18 more per 100 (from 4 more to 36 more) 18 more	⊕⊕⊕○	MODERATE

									44.4%	1.81	per 100 (from 4 more to 36 more)		
Patients reporting side effects - APs: dysthymia only													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁴	none			48/108 (44.4%)	RR 1.23 (0.94 to 1.62)	10 more per 100 (from 3 fewer to 28 more)	⊕⊕⊕⊕ MODERATE	
							57/104 (54.8%)	44.4%	10 more per 100 (from 3 fewer to 28 more)				

¹ Significant heterogeneity - random effects model used

² Inconclusive effect size

³ As 2

⁴ As 2

⁵ Inconclusive effect size; single study

⁶ Single study; non significant effect size

⁷ Non significant effect size

⁸ As 1

⁹ As 2

¹⁰ Single study

¹¹ As 5

¹² As 6

¹³ As 9

¹⁴ As 5

Author(s): NCCMH

Date: 2009-07-13

Question: Should drugs (compared with other drugs) (efficacy data) be used for subthreshold depression?

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Bibliography:

Quality assessment							Summary of findings				Quality	Importance	
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients (compared with other drugs) (efficacy data)	control	Relative (95% CI)	Absolute			
Number of people not achieving at least 50% reduction in depression score: SSRI - Dysthymia => 50% (fluvoxamine vs maprotiline)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	18/24 (75%)	18/24 (75%)	RR 1 (0.72 to 1.39)	0 fewer per 100 (from 21 fewer to 29 more)	⊕⊕⊕⊕ LOW		
								75%		0 fewer per 100 (from 21 fewer to 29 more)			
Number of people not achieving at least 50% reduction in depression score: SSRI - Dysthymia => 50% (SSRI vs amisulpride)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	89/295 (30.2%)	65/299 (21.7%)	RR 1.39 (1.06 to 1.83)	8 more per 100 (from 1 more to 18 more)	⊕⊕⊕⊕ HIGH		
								22%		9 more per 100 (from 1 more to 18 more)			
Number of people not achieving at least 50% reduction in depression score: SSRI - Minor depression only (paroxetine vs maprotiline)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	30/126 (23.8%)	39/119 (32.8%)	RR 0.73 (0.48 to 1.09)	9 fewer per 100 (from 17 fewer to 3 more)	⊕⊕⊕⊕ MODERATE		
								32.8%		9 fewer per 100 (from 17 fewer to 3 more)			
Number of people not achieving at least 50% reduction in depression score: TCA - Dysthymia only (imipramine vs minaprine or moclobemide)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ³	none	46/102 (45.1%)	43/103 (41.7%)	RR 1.07 (0.79 to 1.46)	3 more per 100 (from 9 fewer to 19 more)	⊕⊕⊕⊕ LOW		
								45.1%		3 more per 100 (from 9 fewer to 21 more)			
Number of people not achieving at least 50% reduction in depression score: TCA - Dysthymia => 50% (amitriptyline vs amisulpride)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	34/87 (39.1%)	67/166 (40.4%)	RR 0.97 (0.7 to 1.33)	1 fewer per 100 (from 12 fewer to 13 more)	⊕⊕⊕⊕ LOW		
								40.4%		1 fewer per 100 (from 12 fewer to 13 more)			
Number of people not achieving remission: SSRI - Dysthymia only (sertraline vs imipramine)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	71/134 (53%)	83/136 (61%)	RR 0.87 (0.7 to 1.07)	8 fewer per 100 (from 18 fewer to 4 more)	⊕⊕⊕⊕ MODERATE		
								61%		8 fewer per 100 (from 18 fewer to 4 more)			

											more)		
Number of people not achieving remission: SSRI - Dysthymia => 50% (sertraline vs amisulpride)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	54/156 (34.6%)	42/157 (26.8%)	RR 1.29 (0.92 to 1.81)	8 more per 100 (from 2 fewer to 22 more)	⊕⊕⊕⊕ MODERATE		
								26.8%		8 more per 100 (from 2 fewer to 22 more)			
Number of people not achieving remission: SSRI - Minor depression and subsyndromal depressive symptomatology (49% v 51%) (sertraline vs citalopram)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	42/72 (58.3%)	31/66 (47%)	RR 1.24 (0.9 to 1.71)	11 more per 100 (from 5 fewer to 33 more)	⊕⊕⊕⊕ MODERATE		
								47%		11 more per 100 (from 5 fewer to 33 more)			
Number of people not achieving remission: TCA - Dysthymia only (imipramine vs moclobemide)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	37/68 (54.4%)	34/70 (48.6%)	RR 1.12 (0.81 to 1.55)	6 more per 100 (from 9 fewer to 27 more)	⊕⊕⊕⊕ MODERATE		
								48.6%		6 more per 100 (from 9 fewer to 27 more)			
Mean endpoint scores (clinician rated): SSRI - Dysthymia => 50% (fluvoxamine vs maprotiline) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	21	21	-	SMD 0.01 lower (0.62 lower to 0.59 higher)	⊕⊕⊕⊕ LOW		
Mean endpoint scores (clinician rated): SSRI - Dysthymia => 50% (sertraline vs amisulpride) (Better indicated by lower values)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	279	295	-	SMD 0.16 higher (0 to 0.32 higher)	⊕⊕⊕⊕ HIGH		
Mean endpoint scores (clinician rated): TCA - Dysthymia only (imipramine vs minaprine) (Better indicated by lower values)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ³	none	40	43	-	SMD 0.34 higher (0.1 lower to 0.77 higher)	⊕⊕⊕⊕ LOW		
Mean endpoint scores (clinician rated): TCA - Dysthymia only (amitriptyline vs amisulpride) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	107	101	-	SMD 0.04 higher (0.23 lower to 0.31 higher)	⊕⊕⊕⊕ MODERATE		
Mean endpoint scores (clinician rated): TCA - Dysthymia => 50% (imipramine vs phenelzine) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	16	16	-	SMD 0.73 higher (0.01 to 1.45 higher)	⊕⊕⊕⊕ MODERATE		
Mean endpoint scores (clinician rated): TCA - Dysthymia => 50% (amitriptyline vs amisulpride) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	85	165	-	SMD 0.01 lower (0.27 lower to 0.25 higher)	⊕⊕⊕⊕ MODERATE		
Mean endpoint scores (clinician rated): AP - Dysthymia only (flupenthixol vs ritanserlin) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	36	31	-	SMD 0.26 lower (0.74 lower to 0.22 higher)	⊕⊕⊕⊕ LOW		
Mean change (clinician rated): SSRI - Dysthymia only (sertraline vs imipramine) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	134	136	-	SMD 0.05 higher (0.19 lower to 0.29 higher)	⊕⊕⊕⊕ MODERATE		
Mean change (clinician rated): TCA - Dysthymia only (imipramine vs moclobemide) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	63	67	-	SMD 0.12 higher (0.23 lower to 0.46 higher)	⊕⊕⊕⊕ MODERATE		
Mean change (clinician rated): TCA - Dysthymia only (amitriptyline vs amisulpride) (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	107	101	-	SMD 0.06 higher (0.22 lower to 0.33 higher)	⊕⊕⊕⊕ MODERATE		

¹ Inconclusive effect size; single study

² Single study

³ Inconclusive effect size

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Question: Should drugs compared with others drugs (acceptability/tolerability data) be used for subthreshold depression?

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Quality assessment							Summary of findings				Quality	Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect			
							drugs compared with others drugs (acceptability/tolerability data)	control	Relative (95% CI)	Absolute		
Leaving the study early: SSRI - Dysthymia only (sertraline vs imipramine)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none		45/136		18 fewer per 100		

										(33.1%)		(from 8 fewer to 23 fewer)		
										21/134 (15.7%)	RR 0.47 (0.3 to 0.75)	18 fewer per 100 (from 8 fewer to 23 fewer)	⊕⊕⊕⊕	MODERATE
										33.1%				
Leaving the study early: SSRI - Dysthymia => 50% (fluvoxamine vs maprotiline)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ²	none				4/24 (16.7%)	RR 0.67 (0.22 to 2.07)	8 fewer per 100 (from 19 fewer to 27 more)	⊕⊕⊕⊕	LOW
										6/24 (25%)		8 fewer per 100 (from 19 fewer to 27 more)		
										25%		8 fewer per 100 (from 19 fewer to 27 more)		
Leaving the study early: SSRI - Dysthymia => 50% (sertraline vs amisulpride)														
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ³	none ⁴				67/295 (22.7%)	RR 1.36 (0.98 to 1.89)	6 more per 100 (from 0 fewer to 15 more)	⊕⊕⊕⊕	MODERATE
										50/299 (16.7%)		6 more per 100 (from 0 fewer to 15 more)		
										17%		6 more per 100 (from 0 fewer to 15 more)		
Leaving the study early: SSRI - Minor depression and subsyndromal depressive symptomatology (49% v 51%) (sertraline vs citalopram)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁵	none				20/72 (27.8%)	RR 1.02 (0.59 to 1.75)	5 more per 1000 (from 112 fewer to 205 more)	⊕⊕⊕⊕	LOW
										27.3%				
Leaving the study early: TCA - Dysthymia only (imipramine vs moclobemide)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁶	none				15/103 (14.6%)	RR 1.21 (0.61 to 2.42)	3 more per 100 (from 5 fewer to 17 more)	⊕⊕⊕⊕	LOW
										13/108 (12%)		3 more per 100 (from 5 fewer to 17 more)		
										12%		3 more per 100 (from 5 fewer to 17 more)		
Leaving the study early: TCA - Dysthymia only (amitriptyline vs amisulpride)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁷	none				20/111 (18%)	RR 1.34 (0.71 to 2.51)	5 more per 100 (from 4 fewer to 20 more)	⊕⊕⊕⊕	LOW
										14/104 (13.5%)		5 more per 100 (from 4 fewer to 20 more)		
										13.5%		5 more per 100 (from 4 fewer to 20 more)		
Leaving the study early: TCA - Dysthymia => 50% (imipramine vs phenelzine)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁸	none				5/37 (13.5%)	RR 1.22 (0.35 to 4.17)	2 more per 100 (from 7 fewer to 35 more)	⊕⊕⊕⊕	LOW
										4/36 (11.1%)		2 more per 100 (from 7 fewer to 35 more)		
										11.1%		2 more per 100 (from 7 fewer to 35 more)		
Leaving the study early: TCA - Dysthymia => 50% (amitriptyline vs amisulpride)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁹	none				41/87 (47.1%)	RR 1.07 (0.81 to 1.42)	3 more per 100 (from 8 fewer to 18 more)	⊕⊕⊕⊕	LOW
										73/166 (44%)		3 more per 100 (from 8 fewer to 18 more)		
										44%		3 more per 100 (from 8 fewer to 18 more)		
Leaving the study early: AP (flupenthixol vs ritanserin)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹⁰	none				3/36 (8.3%)	RR 1.29 (0.23 to 7.24)	2 more per 100 (from 5 fewer to 40 more)	⊕⊕⊕⊕	LOW
										2/31 (6.5%)		2 more per 100 (from 5 fewer to 40 more)		
										6.5%		2 more per 100 (from 5 fewer to 41 more)		
Leaving the study early due to side effects: SSRI - Dysthymia only (sertraline vs imipramine)														
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹¹	none				8/134 (6%)	RR 0.32 (0.15 to 0.70)	12 fewer per 100 (from 6 fewer to 16 fewer)	⊕⊕⊕⊕	MODERATE
										25/136 (18.4%)		13 fewer		

									18.4%	0.69	per 100 (from 6 fewer to 16 fewer)		
Leaving the study early due to side effects: SSRI - Dysthymia => 50% (sertraline/fluoxetine vs amisulpride)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹²	none		22/295 (7.5%)	23/299 (7.7%)	RR 0.97 (0.55 to 1.7)	0 fewer per 100 (from 3 fewer to 5 more)	⊕⊕⊕⊕ LOW	
								7.8%			0 fewer per 100 (from 4 fewer to 5 more)		
Leaving the study early due to side effects: SSRI - Minor depression and subsyndromal depressive symptomatology (49% v 51%) (sertraline vs citalopram)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹³	none		8/72 (11.1%)	10/66 (15.2%)	RR 0.73 (0.31 to 1.75)	4 fewer per 100 (from 10 fewer to 11 more)	⊕⊕⊕⊕ LOW	
								15.2%			4 fewer per 100 (from 10 fewer to 11 more)		
Leaving the study early due to side effects: TCA - Dysthymia only (imipramine vs minaprine/moclobemide)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹⁴	none		15/137 (10.9%)	10/141 (7.1%)	RR 1.54 (0.72 to 3.3)	4 more per 100 (from 2 fewer to 16 more)	⊕⊕⊕⊕ LOW	
								7.8%			4 more per 100 (from 2 fewer to 18 more)		
Leaving the study early due to side effects: TCA - Dysthymia only (amitriptyline vs amisulpride)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹⁵	none		6/111 (5.4%)	3/104 (2.9%)	RR 1.87 (0.48 to 7.3)	3 more per 100 (from 1 fewer to 18 more)	⊕⊕⊕⊕ LOW	
								2.9%			3 more per 100 (from 2 fewer to 18 more)		
Leaving the study early due to side effects: TCA - Dysthymia => 50% (amitriptyline vs amisulpride)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹⁶	none		11/87 (12.6%)	23/166 (13.9%)	RR 0.91 (0.47 to 1.78)	1 fewer per 100 (from 7 fewer to 11 more)	⊕⊕⊕⊕ LOW	
								13.9%			1 fewer per 100 (from 7 fewer to 11 more)		
Leaving the study early due to side effects: AP - Dythymia only (flupenthixol vs ritanserin)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹⁷	none		2/36 (5.6%)	2/33 (6.1%)	RR 0.92 (0.14 to 6.14)	0 fewer per 100 (from 5 fewer to 31 more)	⊕⊕⊕⊕ LOW	
								6.1%			0 fewer per 100 (from 5 fewer to 31 more)		
Patients reporting side effects: SSRI - Dysthymia => 50% (sertraline vs amisulpride)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none		130/293 (44.4%)	137/298 (46%)	RR 0.96 (0.81 to 1.15)	2 fewer per 100 (from 9 fewer to 7 more)	⊕⊕⊕⊕ HIGH	
								46.1%			2 fewer per 100 (from 9 fewer to 7 more)		
Patients reporting side effects: TCA - Dysthymia only (imipramine vs minaprine)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁸	none		20/34 (58.8%)	14/33 (42.4%)	RR 1.39 (0.85 to 2.26)	17 more per 100 (from 6 fewer to 53 more)	⊕⊕⊕⊕ MODERATE	
								42.4%			17 more per 100 (from 6 fewer to 53 more)		
Patients reporting side effects: TCA - Dysthymia only (amitriptyline vs amisulpride)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹⁹	none		69/111 (62.2%)	57/104 (54.8%)	RR 1.13 (0.9 to 1.4)	7 more per 100 (from 5 fewer to 23 more)	⊕⊕⊕⊕ MODERATE	
											7 more		

								54.8%	1.42	per 100 (from 5 fewer to 23 more)		
Patients reporting side effects: TCA - Dysthymia => 50% (amitriptyline vs amisulpride)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²⁰	none	62/85 (72.9%)	106/165 (64.2%)	RR 1.14 (0.96 to 1.35)	9 more per 100 (from 3 fewer to 22 more)	⊕⊕⊕○ MODERATE	
								64.2%				9 more per 100 (from 3 fewer to 22 more)
Patients reporting side effects: AP - Dysthymia only (flupenthixol vs ritanserlin)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ²¹	none	16/36 (44.4%)	15/33 (45.5%)	RR 0.98 (0.58 to 1.65)	1 fewer per 100 (from 19 fewer to 30 more)	⊕⊕○○ LOW	
								45.5%				1 fewer per 100 (from 19 fewer to 30 more)

- ¹ Single study
- ² Inconclusive effect size; single study
- ³ No explanation was provided
- ⁴
- ⁵ As 2
- ⁶ As 2
- ⁷ As 2
- ⁸ As 2
- ⁹ As 2
- ¹⁰ As 2
- ¹¹ As 1
- ¹² As 2
- ¹³ As 2
- ¹⁴ Inconclusive effect size
- ¹⁵ As 2
- ¹⁶ As 2
- ¹⁷ As 2
- ¹⁸ As 1
- ¹⁹ As 1
- ²⁰ As 1
- ²¹ As 2

Author(s): NCCMH
Date: 2008-12-02
Question: Should Relapse prevention be used for dysthymia?
Settings:
Bibliography:

Quality assessment							Summary of findings				Quality	Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect			
							Relapse prevention	control	Relative (95% CI)	Absolute		
Recurrence - TCAs												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	0/14 (0%)	6/13 (46.2%)	RR 0.07 (0 to 1.16)	43 fewer per 100 (from 46 fewer to 7 more)	⊕⊕○○ LOW	
								46.2%				43 fewer per 100 (from 46 fewer to 7 more)

- ¹ Inconclusive effect size; single study

Author(s): NCCMH

Date: 2009-07-10

Question: Should Psychological therapies vs no-treatment control be used for depression symptoms which do not meet criteria for major depressive disorder?

Settings:

Bibliography:

Quality assessment							Summary of findings					Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect		Quality	
							Psychological therapies	no-treatment control	Relative (95% CI)	Absolute		
Efficacy data - Number not responding												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	72/139 (51.8%)	84/138 (60.9%) 58.9%	RR 0.86 (0.70 to 1.06)	9 fewer per 100 (from 18 fewer to 4 more) 8 fewer per 100 (from 18 fewer to 4 more)	⊕⊕⊕⊕ MODERATE	
Efficacy data - Number not achieving remission												
1	randomised trials	no serious limitations	serious ²	no serious indirectness	serious ¹	none	62/115 (53.9%)	70/112 (62.5%) 58.8%	RR 0.86 (0.69 to 1.08)	9 fewer per 100 (from 19 fewer to 5 more) 8 fewer per 100 (from 18 fewer to 5 more)	⊕⊕⊕⊕ LOW	
Efficacy data (continuous) - Clinician-rated endpoint scores (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	97	99	-	SMD 0.27 lower (0.55 lower to 0.01 higher)	⊕⊕⊕⊕ MODERATE	
Acceptability and tolerability data - Leaving treatment early for any reason												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	20/139 (14.4%)	23/138 (16.7%) 13.2%	RR 0.86 (0.50 to 1.47)	2 fewer per 100 (from 8 fewer to 8 more) 2 fewer per 100 (from 7 fewer to 6 more)	⊕⊕⊕⊕ MODERATE	

¹ Inconclusive effect² Significant heterogeneity - random effects model used

Author(s): NCCMH

Date: 2009-07-10

Question: Should Psychological therapies vs antidepressants be used for depression symptoms which do not meet criteria for major depressive disorder?

Settings:

Bibliography:

Quality assessment							Summary of findings					Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect		Quality	
							Psychological therapies	antidepressants	Relative (95% CI)	Absolute		
Efficacy data - Number not responding												
3	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	92/162 (56.8%)	81/157 (51.6%) 51.9%	RR 1.09 (0.92 to 1.29)	5 more per 100 (from 4 fewer to 15 more) 5 more per 100 (from 4 fewer to 15 more)	⊕⊕⊕⊕ HIGH	
Efficacy data - Number not remitting												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	80/138 (58%)	70/135 (51.9%) 58.3%	RR 1.14 (0.92 to 1.41)	7 more per 100 (from 4 fewer to 21 more) 8 more per 100 (from 5 fewer to 24 more)	⊕⊕⊕⊕ MODERATE	
Efficacy data (continuous) - Clinician-rated mean endpoint (Better indicated by lower values)												
4	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	308	320	-	SMD 0.29 higher (0.13 to 0.45 higher)	⊕⊕⊕⊕ HIGH	
Efficacy data (continuous) - Clinician-rated mean endpoint 6-month follow-up (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	167	186	-	SMD 0.19 higher (0.02 lower to 0.4 higher)	⊕⊕⊕⊕ MODERATE	
Efficacy data (continuous) - Clinician-rated mean endpoint 18-month follow-up (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ³	none	156	179	-	SMD 0.26 higher (0.05 to 0.48 higher)	⊕⊕⊕⊕ MODERATE	
Efficacy data (continuous) - Self-rated mean endpoint (Better indicated by lower values)												
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	no serious imprecision	none	32	35	-	SMD 0.37 higher (0.11 lower to	⊕⊕⊕⊕ HIGH	

											0.86 higher)		
Acceptability and tolerability data - Leaving treatment early for any reason													
4	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none		39/175 (22.3%)		RR 0.67 (0.42 to 1.06)	7 fewer per 100 (from 13 fewer to 1 more)	⊕⊕⊕○	MODERATE
							25/175 (14.3%)		23%		8 fewer per 100 (from 13 fewer to 1 more)		
Acceptability and tolerability data - Leaving treatment early due to side effects													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ⁴	none		1/18 (5.6%)		RR 0.45 (0.02 to 10.3)	3 fewer per 100 (from 5 fewer to 52 more)	⊕⊕○○	LOW
							0/13 (0%)		5.6%		3 fewer per 100 (from 5 fewer to 52 more)		

¹ Inconclusive effect
² Single study
³ Significant heterogeneity - random effects model used
⁴ Single study; inconclusive effect size

Author(s): NCCMH
Date: 2009-07-10
Question: Should Psychological therapies + antidepressants vs antidepressants be used for depression symptoms which do not meet criteria for major depressive disorder?
Settings:
Bibliography:

Quality assessment							Summary of findings				Quality	Importance	
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect				
							Psychological therapies + antidepressants	antidepressants	Relative (95% CI)	Absolute			
Efficacy data - Number not responding													
2	randomised trials	no serious limitations	serious ¹	no serious indirectness	serious ²	none	28/46 (60.9%)	30/46 (65.2%)		RR 0.96 (0.52 to 1.79)	3 fewer per 100 (from 31 fewer to 52 more)	⊕⊕○○	LOW
								64.4%			3 fewer per 100 (from 31 fewer to 51 more)		
Efficacy data - Number not remitting													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ³	none	10/21 (47.6%)	14/24 (58.3%)		RR 0.82 (0.47 to 1.43)	10 fewer per 100 (from 31 fewer to 25 more)	⊕⊕○○	LOW
								58.3%			10 fewer per 100 (from 31 fewer to 25 more)		
Efficacy data (continuous) - clinician-rated mean endpoint (Better indicated by lower values)													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	233	220	-		SMD 0.09 higher (0.1 lower to 0.27 higher)	⊕⊕⊕○	MODERATE
Efficacy data (continuous) - Clinician-rated mean endpoint at 6-month follow-up (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁴	none	196	186	-		SMD 0.01 higher (0.19 lower to 0.21 higher)	⊕⊕⊕○	MODERATE
Efficacy data (continuous) - Clinician-rated mean endpoint at 18-month follow-up (Better indicated by lower values)													
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ⁴	none	190	179	-		SMD 0.06 higher (0.14 lower to 0.27 higher)	⊕⊕⊕○	MODERATE
Acceptability and tolerability data - Leaving treatment early for any reason													
2	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ²	none	5/46 (10.9%)	5/46 (10.9%)		RR 1.09 (0.37 to 3.25)	1 more per 100 (from 7 fewer to 24 more)	⊕⊕⊕○	MODERATE
								10.4%			1 more per 100 (from 7 fewer to 23 more)		

¹ Significant heterogeneity - random effects model used
² Inconclusive effect
³ Single study; inconclusive effect
⁴ Single study

Author(s): NCCMH

Date: 2009-07-10

Question: Should Psychological therapies + antidepressants vs psychological therapies be used for depression symptoms which do not meet criteria for major depressive disorder?

Settings:

Bibliography:

Quality assessment							Summary of findings					Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect		Quality	
							Psychological therapies + antidepressants	psychological therapies	Relative (95% CI)	Absolute		
Efficacy data - Number not responding												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ¹	none	8/25 (32%)	16/24 (66.7%)	RR 0.48 (0.25 to 0.91)	35 fewer per 100 (from 6 fewer to 50 fewer)	⊕⊕⊕⊕ MODERATE	
								66.7%		35 fewer per 100 (from 6 fewer to 50 fewer)		
Efficacy data (continuous) - Clinician-rated mean endpoint data (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ^{1,2}	none	212	178	-	SMD 0.17 lower (0.37 lower to 0.03 higher)	⊕⊕⊕⊕ MODERATE	
Efficacy data (continuous) - Clinician-rated mean endpoint data 6-month follow-up (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ^{1,2}	none	196	167	-	SMD 0.18 lower (0.38 lower to 0.03 higher)	⊕⊕⊕⊕ MODERATE	
Efficacy data (continuous) - Clinician-rated mean endpoint data 18-month follow-up (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ^{1,2}	none	190	156	-	SMD 0.2 lower (0.41 lower to 0.01 higher)	⊕⊕⊕⊕ MODERATE	
Acceptability and tolerability data - Leaving treatment early for any reason												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	serious ^{1,2}	none	1/25 (4%)	0/24 (0%)	RR 2.88 (0.12 to 67.53)	0 more per 100 (from 0 fewer to 0 more)	⊕⊕⊕⊕ MODERATE	
								0%		0 more per 100 (from 0 fewer to 0 more)		

¹ Single study

² Inconclusive effect

Author(s): NCCMH

Date: 2009-01-22

Question: Should Short Psychodynamic Verbal v Short Psychodynamic Art be used for depression symptoms which do not meet criteria for major depressive disorder?

Settings:

Bibliography:

Quality assessment							Summary of findings					Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect		Quality	
							Short Psychodynamic Verbal v Short Psychodynamic Art	control	Relative (95% CI)	Absolute		
Efficacy data - Self-rated mean endpoint (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	21	18	-	SMD 0.11 lower (0.74 lower to 0.52 higher)	⊕⊕⊕⊕ LOW	
Efficacy data - Self-rated mean endpoint at 3-month follow-up (Better indicated by lower values)												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	21	18	-	SMD 0.26 lower (0.9 lower to 0.37 higher)	⊕⊕⊕⊕ LOW	
Acceptability and tolerability data - Leaving treatment early for any reason												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	1/22 (4.5%)	3/21 (14.3%)	RR 0.32 (0.04 to 2.82)	10 fewer per 100 (from 14 fewer to 26 more)	⊕⊕⊕⊕ LOW	
								14.3%		10 fewer per 100 (from 14 fewer to 26 more)		

¹ Single study; inconclusive effect

Author(s): NCCMH

Date: 2009-01-22

Question: Should Partial responders be used for depression symptoms which do not meet criteria for major depressive disorder?

Settings:

Bibliography:

Quality assessment							Summary of findings				Quality	Importance
No of studies	Design	Limitations	Inconsistency	Indirectness	Imprecision	Other considerations	No of patients		Effect			
							Partial responders	control	Relative (95% CI)	Absolute		
Number of people not achieving at least 50% reduction in depression score - Psych/SSRI Combo v SSRI												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	4/20 (20%)	7/20 (35%)	RR 0.57 (0.2 to 1.65)	15 fewer per 100 (from 28 fewer to 23 more)	⊕⊕○○ LOW	
								35%		15 fewer per 100 (from 28 fewer to 23 more)		
Number of people not achieving at least 50% reduction in depression score: 12 week follow-up - Psych/SSRI Combo v SSRI												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	9/20 (45%)	14/20 (70%)	RR 0.64 (0.37 to 1.13)	25 fewer per 100 (from 44 fewer to 9 more)	⊕⊕○○ LOW	
								70%		25 fewer per 100 (from 44 fewer to 9 more)		
Number of people not achieving remission - Psych/SSRI Combo v SSRI												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	6/20 (30%)	10/20 (50%)	RR 0.6 (0.27 to 1.34)	20 fewer per 100 (from 37 fewer to 17 more)	⊕⊕○○ LOW	
								50%		20 fewer per 100 (from 37 fewer to 17 more)		
Number of people not achieving remission: 12 week follow-up - Psych/SSRI Combo v SSRI												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	16/20 (80%)	14/20 (70%)	RR 1.14 (0.8 to 1.64)	10 more per 100 (from 14 fewer to 45 more)	⊕⊕○○ LOW	
								70%		10 more per 100 (from 14 fewer to 45 more)		
Leaving the study early - Psych/SSRI Combo v SSRI												
1	randomised trials	no serious limitations	no serious inconsistency	no serious indirectness	very serious ¹	none	2/20 (10%)	3/20 (15%)	RR 0.67 (0.12 to 3.57)	5 fewer per 100 (from 13 fewer to 39 more)	⊕⊕○○ LOW	
								15%		5 fewer per 100 (from 13 fewer to 39 more)		

¹ Single study; inconclusive effect