GRADE PROFILES FOR ECONOMIC EVIDENCE

Clinical / economic question: escitalopram versus placebo

Escitalopram versus placebo Increment Incremental **ICER** Study & Limitations **Applicability** Other comments al cost Uncertainty country effect (£/effect) (£)¹ Guideline • Time horizon: 42 weeks analysis Directly Not relevant; both interventions dominated by sertraline; probability of Minor Model included 6 drugs Escitalopram -£84.70 0.0395 limitations² applicable³ UK plus no treatment dominant sertraline being cost-effective at £20,000/QALY: 0.71 (placebo)

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: sertraline versus placebo

Sertrali	Sertraline versus placebo									
Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty			
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo) Time horizon: 42 weeks	-£168.383	0.0422	Sertraline dominant	Probability of sertraline being cost-effective at £20,000/QALY: 0.71			

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: paroxetine versus placebo

Directly

applicable³

Paroxetine versus placebo Study & country Limitations Applicability Other comments Increment al cost (£)¹ Incremental effect (£/effect) Uncertainty

0.0363

1. Costs expressed in 2009 UK pounds

Minor

limitations²

analysis

UK

2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered

Paroxetine

dominant

Not relevant; both interventions dominated by sertraline; probability of

sertraline being cost-effective at £20,000/QALY: 0.71

3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

-£126.176

Model included 6 drugs

plus no treatment

(placebo)

Clinical / economic question: duloxetine versus placebo

Duloxe	Duloxetine versus placebo									
Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty			
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo) Time horizon: 42 weeks	-£35.096	0.0404	Duloxetine dominant	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71			

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: venlafaxine XL versus placebo

Venlafaxine XL versus placebo

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	-£109.156	0.0399	Venlafaxine XL dominant	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: pregabalin versus placebo

Pregabalin versus placebo

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	-£13.178	0.0420	Pregabalin dominant	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: escitalopram versus paroxetine

Escitalopram versus paroxetine

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect) ¹	Uncertainty ¹
Iskedjian <i>et</i> <i>al.</i> , 2008 Canada	Potentially serious limitations ²	Partially applicable ³	Measure of outcome: number of symptom-free days (SFDs) Time horizon: 24 weeks	£32	9.4SFDs	£3.4/SFD	£2.9-£4.49/SFD
Jørgensen et al., 2006 UK	Potentially serious limitations ⁴	Directly applicable ⁵	Measure of outcome: % of people with maintained response Time horizon: 36 weeks	-£45	7.7% more people with maintained response	Escitalopram dominant	Escitalopram dominant
Guideline analysis UK	Minor limitations ⁶	Directly applicable ⁷	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	£41.5	0.0032	£12,893/ QALY	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs converted and uplifted to 2009 UK pounds, using PPP exchange rates (http://www.oecd.org/std/ppp) and the UK HCHS inflation index.
- 2. Efficacy data derived selectively from one RCT; many clinical and all resource use estimates based on expert opinion; limited sensitivity analysis; funded by industry
- 3. Conducted in Canada Ministry of Health perspective (direct healthcare costs considered); no QALYs estimated but outcome measure considered relevant; utility scores for GAD are still scarce and of low quality
- 4. Efficacy data derived selectively from one RCT; some clinical and resource use estimates based on expert opinion; limited sensitivity analysis; funded by industry
- 5. NHS perspective; no QALYs estimated but outcome measure considered relevant; utility scores for GAD are still scarce and of low quality
- 6. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 7. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: sertraline versus paroxetine

Sertraline versus paroxetine

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	-£42.207	0.0059	Sertraline dominant	Probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: escitalopram versus venlafaxine XL

Escitalopram versus venlafaxine XL

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	£24.458	-0.0004	Venlafaxine XL dominant	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: duloxetine versus venlafaxine XL

Duloxetine versus venlafaxine XL

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	£74.060	0.0005	£150,761 /QALY	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: venlafaxine XL versus pregabalin

Venlafaxine XL versus pregabalin

Study & country	Limitations	Applicability	Other comments	Increment al cost (£) ¹	Incremental effect	ICER (£/effect)	Uncertainty
Guideline analysis UK	Minor limitations ²	Directly applicable ³	Time horizon: 42 weeks Model included 6 drugs plus no treatment (placebo)	£17.020	0.0036	£4,771 /QALY	Not relevant; both interventions dominated by sertraline; probability of sertraline being cost-effective at £20,000/QALY: 0.71

- 1. Costs expressed in 2009 UK pounds
- 2. Evidence synthesis based on network (mixed treatment comparison) meta-analytic techniques; resource use based on data reported in RCTs, a national survey and GDG expert opinion; impact of tolerable side effects on health-related quality of life not considered; costs associated with management of side effects no considered
- 3. Analysis conducted to assist guideline development; NHS & personal social services perspective; QALYs estimated based on SF-6D

Clinical / economic question: venlafaxine XL versus diazepam

Venlafaxine XL versus diazepam

Study & country	Limitations	Applicability	Other comments	Incrementa I cost (£) ¹	Incremental effect	ICER (£/effect) ¹	Uncertainty ¹
Guest et al., 2004 UK	Potentially serious limitations ²	Partially applicable ³	Measure of outcome: percentage of people with successful treatment defined as CGI score of 1 at 6 months Time horizon: 6 months	£56	10.8% extra successfully treated people	£516/ successfully treated person	Venlafaxine XL dominates - £2,203/successfully treated person Probabilistic analysis: venlafaxine XL dominated diazepam in at least 25% of iterations

- 1. Costs uplifted to 2009 UK pounds using the UK HCHS inflation index.
- 2. Efficacy data derived selectively from one RCT; resource use estimated based on expert opinion; limited sensitivity analysis; funded by industry
- 3. UK / NHS perspective; no QALYs estimated but outcome measure considered relevant; utility scores for GAD are still scarce and of low quality

Clinical / economic question: paroxetine versus venlafaxine XL

Paroxetine versus venlafaxine XL

Study & country	Limitations	Applicability	Other comments	Incrementa I cost (£) ¹	Incremental effect	ICER (£/effect) ¹	Uncertainty ¹
Vera- Llonch et al., 2010 Spain	Potentially serious limitations ²	Partially applicable ³	Time horizon: 12 months, but treatment effect assumed to last from 8 weeks (end of treatment) until 12 months	£468	0.027	£17,565/ QALY	£14,567-£26,442/QALY Probabilistic analysis: pregabalin cost effective in roughly 95% of iterations at a cost effectiveness threshold of £20,000/QALY

- 1. Costs converted and uplifted to 2009 UK pounds, using PPP exchange rates (http://www.oecd.org/std/ppp) and the UK HCHS inflation index.
- 2. Efficacy data derived selectively from one RCT; treatment effect assumed to last for 44 weeks beyond end of treatment; funded by industry
- 3. Spanish third party payer perspective; valuation of QALYs derived from Spanish population