Appendix 16b: Forest plots for clinical evidence reviews on pharmacological interventions

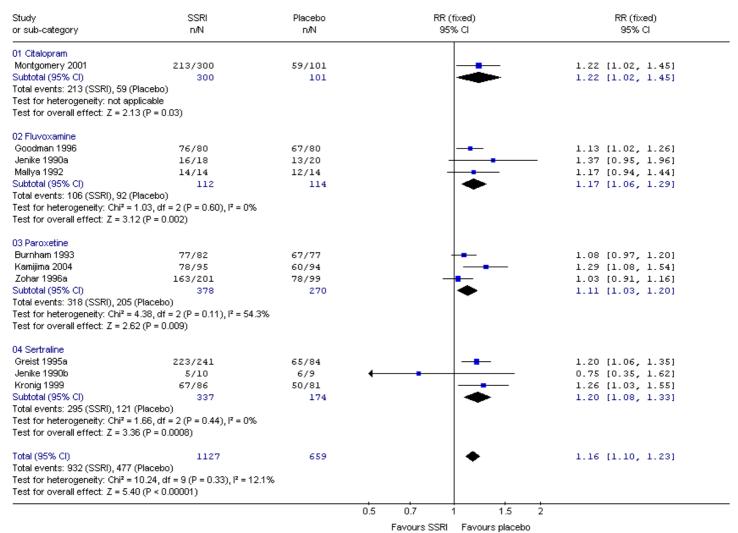
SSRIs (OCD, adults); Clomipramine (OCD, adults); TCAs (OCD); SNRIs (OCD); MAOIs (OCD); Anxiolytics (OCD); Other Pharmacological (OCD); Augmentation (OCD); SSRIs (OCD, children); Clomipramine (OCD, children); Pharmacological (BDD)

SSRIs (OCD, adults)

Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)

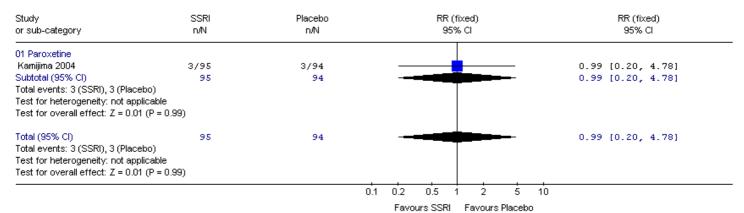
Outcome: 01 Adverse events



Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)

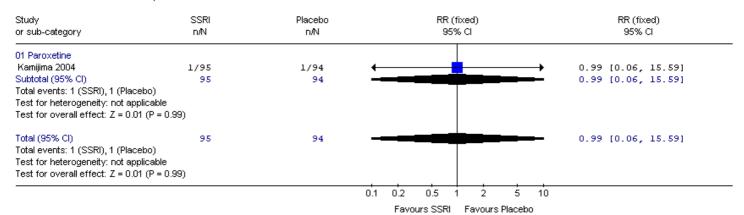
Outcome: 02 Serious adverse events



Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)

Outcome: 03 Suicide attempts



Review: OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase) Outcome: 04 Leaving study early SSRI Placebo RR (fixed) RR (fixed) Study or sub-category n/Ν n/N 95% CI 95% CI 01 Citalopram Montgomery 2001 46/300 17/101 0.91 [0.55, 1.51] Subtotal (95% CI) 300 101 0.91 [0.55, 1.51] Total events: 46 (SSRI), 17 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.36 (P = 0.72) 02 Fluoxetine Jenike 1997 4/23 3/21 1.22 [0.31, 4.82] Montgomery 1993 41/160 15/57 0.97 [0.59, 1.62] Subtotal (95% CI) 1.00 [0.62, 1.62] 183 78 Total events: 45 (SSRI), 18 (Placebo) Test for heterogeneity: $Chi^2 = 0.09$, df = 1 (P = 0.76), $I^2 = 0\%$ Test for overall effect: Z = 0.02 (P = 0.99) 03 Fluvoxamine Goodman 1989 2/23 6/23 0.33 [0.07, 1.48] Goodman 1996 23/80 17/80 1.35 [0.78, 2.33] 1.38 [0.93, 2.03] Hollander 2003b 43/127 31/126 Jenike 1990a 0/18 0/20 Not estimable 0.88 [0.32, 2.40] 1.00 [0.07, 13.87] Mallya 1992 5/19 6/20 Perse 1987 1/10 1/10 Subtotal (95% CI) 279 1.21 [0.91, 1.62] Total events: 74 (SSRI), 61 (Placebo) Test for heterogeneity: $Chi^2 = 3.85$, df = 4 (P = 0.43), $I^2 = 0\%$ Test for overall effect: Z = 1.30 (P = 0.19) 04 Paroxetine Burnham 1993 20/77 1.31 [0.81, 2.13] 28/82 1.21 [0.72, 2.04] Hollander 2003d 53/259 15/89 Zohar 1996a 0.62 [0.44, 0.87] 49/201 39/99 Subtotal (95% CI) 542 265 0.91 [0.71, 1.16] Total events: 130 (SSRI), 74 (Placebo) Test for heterogeneity: $Chi^2 = 8.21$, df = 2 (P = 0.02), $I^2 = 75.6\%$ Test for overall effect: Z = 0.76 (P = 0.45) 05 Sertraline Chouinard 1990 0.44 [0.12, 1.59] 3/43 7/44 Greist 1995a 65/241 24/84 0.94 [0.63, 1.40] Jenike 1990b 0/10 0/9 Not estimable Kronig 1999 25/86 25/81 0.94 [0.59, 1.50] Subtotal (95% CI) 218 0.89 [0.67, 1.20] 380 Total events: 93 (SSRI), 56 (Placebo) Test for heterogeneity: $Chi^2 = 1.30$, df = 2 (P = 0.52), $I^2 = 0\%$ Test for overall effect: Z = 0.76 (P = 0.44) Total (95% CI) 941 0.98 [0.85, 1.13]

> 0.1 0.2

0.5

Favours SSRI Favours placebo

5 10

1682

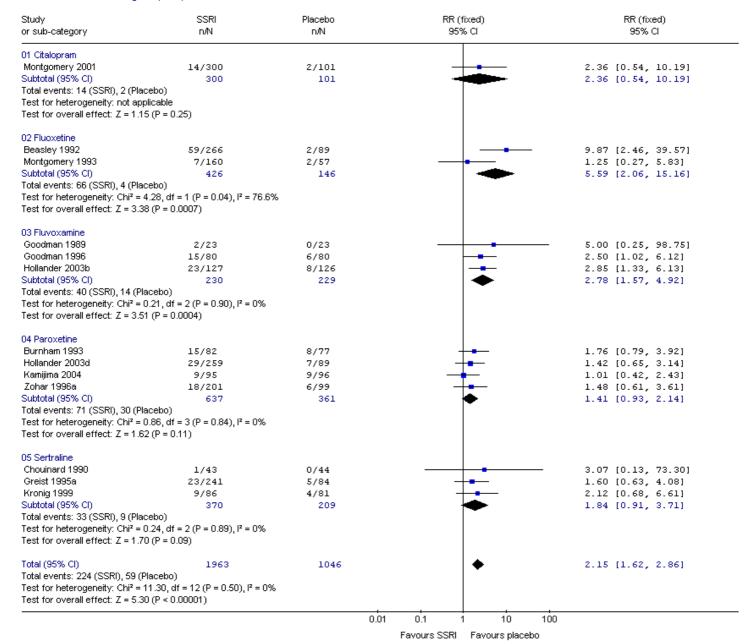
Total events: 388 (SSRI), 226 (Placebo)

Test for overall effect: Z = 0.26 (P = 0.79)

Test for heterogeneity: Chi² = 16.94, df = 13 (P = 0.20), l² = 23.3%

Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)
Outcome: 05 Leaving study early due to adverse events



OCD adults: SSRIs

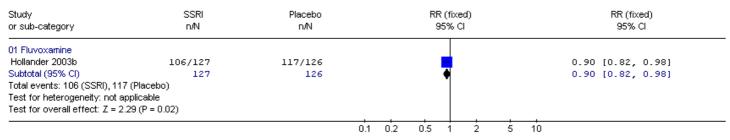
Review:

Comparison: 01 SSRIs vs placebo (acute phase) 06 Non-responders (OCD) (CGI or Y-BOCS) Outcome: SSRI Placebo RR (fixed) RR (fixed) Study 95% CI or sub-category n/Ν n/N 95% CI 01 Citalopram Montgomery 2001 126/300 64/101 0.66 [0.54, 0.81] Subtotal (95% CI) 300 101 0.66 [0.54, 0.81] Total events: 126 (SSRI), 64 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 4.05 (P < 0.0001)02 Fluoxetine Beasley 1992 190/266 82/89 0.78 [0.70, 0.85] Montgomery 1993 46/57 0.71 [0.59, 0.86] 92/160 0.75 [0.69, 0.82] Subtotal (95% CI) Total events: 282 (SSRI), 128 (Placebo) Test for heterogeneity: Chi² = 0.69, df = 1 (P = 0.41), l² = 0% Test for overall effect: Z = 6.18 (P < 0.00001) 03 Fluvoxamine Hollander 2003b 74/127 90/126 0.82 [0.68, 0.98] 0.72 [0.52, 0.99] Mallya 1992 13/19 19/20 0.80 [0.68, 0.94] Subtotal (95% CI) 146 146 Total events: 87 (SSRI), 109 (Placebo) Test for heterogeneity: $Chi^2 = 0.45$, df = 1 (P = 0.50), $I^2 = 0\%$ Test for overall effect: Z = 2.69 (P = 0.007) 04 Paroxetine 0.98 [0.75, 1.28] Burnham 1993 47/82 45/77 0.85 [0.73, 1.00] Hollander 2003d 161/259 65/89 Kamiiima 2004 48/95 74/96 0.66 [0.52, 0.82] Zohar 1996a 92/201 64/99 0.71 [0.57, 0.87] Subtotal (95% CI) 0.78 [0.71, 0.87] 361 Total events: 348 (SSRI), 248 (Placebo) Test for heterogeneity: $Chi^2 = 7.09$, df = 3 (P = 0.07), $I^2 = 57.7\%$ Test for overall effect: Z = 4.66 (P < 0.00001) 05 Sertraline Greist 1995a 148/241 0.87 [0.74, 1.04] 59/84 Subtotal (95% CI) 0.87 [0.74, 1.04] 241 84 Total events: 148 (SSRI), 59 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.54 (P = 0.12) Total (95% CI) 1750 838 0.77 [0.73, 0.82] Total events: 991 (SSRI), 608 (Placebo) Test for heterogeneity: $Chi^2 = 12.76$, df = 9 (P = 0.17), $I^2 = 29.5\%$ Test for overall effect: Z = 8.51 (P < 0.00001)0.5 0.715 Favours SSRI Favours placebo OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase) 07 Non-responders (MDD) Outcome: Study SSRI Placebo RR (fixed) RR (fixed) or sub-category n/N n/N 95% CI 95% CI 01 Fluvoxamine Goodman 1989 18/23 20/23 0.90 [0.69, 1.18] Subtotal (95% CI) 23 23 0.90 [0.69, 1.18] Total events: 18 (SSRI), 20 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.77 (P = 0.44) 0.5 10 0.2 Favours SSRI Favours placebo

Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)

Outcome: 08 Non-remitters



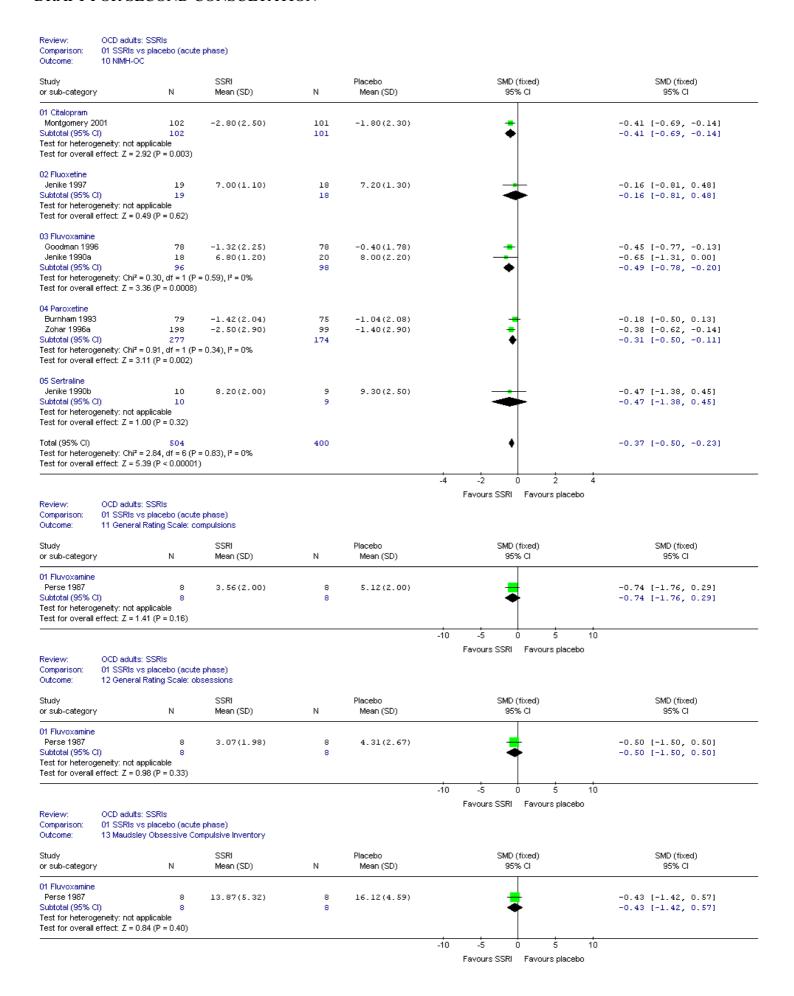
Favours SSRI Favours placebo

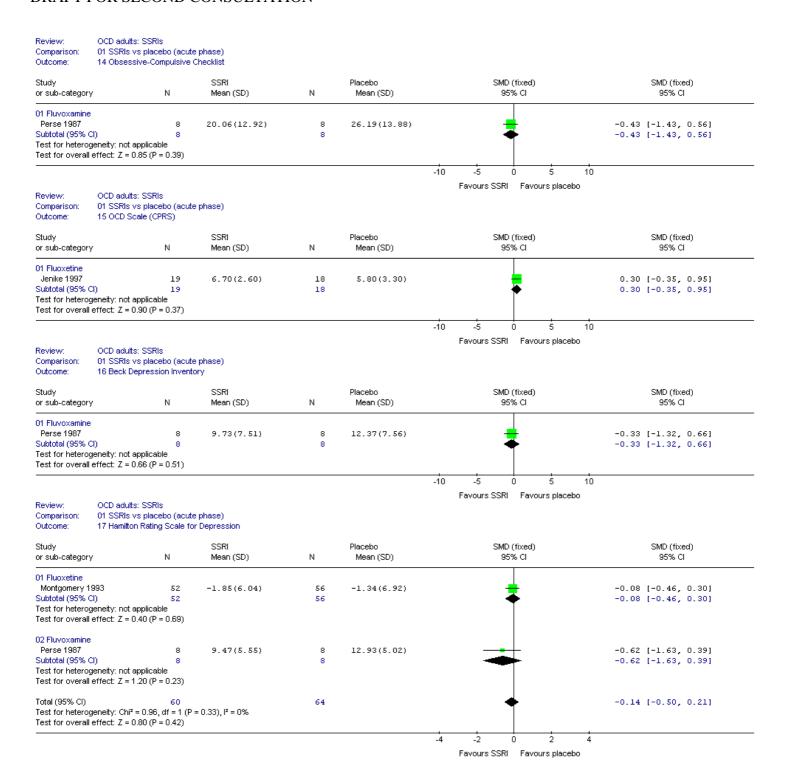
Review: OCD adults: SSRIs

Comparison: 01 SSRIs vs placebo (acute phase)

Outcome: 09 Y-BOCS

Study or sub-category N		SRI ean (SD)	N	Placebo Mean (SD)	SMD (* 95%		SMD (fixed) 95% Cl
01 Citalopram							
Montgomery 2001 10:		40(7.30)	101	-5.60(6.90)	-		-0.39 [-0.67, -0.11]
Subtotal (95% CI) 103	2		101		•		-0.39 [-0.67, -0.11]
est for heterogeneity: not applicable est for overall effect: Z = 2.77 (P = 0.0)	106)						
2 Fluoxetine							
Beasley 1992 7:	E 10	.90(8.30)	76	23.60(7.50)			-0.59 [-0.92, -0.27]
Jenike 1997		.20(6.30)	18	18.70(6.10)			-0.39 [-1.05, 0.26]
Montgomery 1993 52		.13(6.41)	56	-3.70(5.98)			-0.23 [-0.61, 0.15]
ubtotal (95% CI) 140		10,0.11,	150	0.70,0.50,	•		-0.43 [-0.66, -0.20]
est for heterogeneity: Chi² = 2.03, df =		. I² = 1.5%	200		•		0.10 (0.00, 0.20)
est for overall effect: Z = 3.66 (P = 0.0		,, ,,,,,,					
13 Fluvoxamine							
Goodman 1989 2:	1 19.	40(7.00)	21	28.00(7.00)			-1.21 [-1.87, -0.54]
Goodman 1996 78	8 -3.	.95(6.28)	78	-1.71(4.88)	-		-0.40 [-0.71, -0.08]
Hollander 2003b 111	7 -8.	.50(7.57)	120	-5.60(7.67)	-		-0.38 [-0.64, -0.12]
Jenike 1990a 18	8 18.	.80(4.00)	20	21.80(7.60)	-		-0.48 [-1.12, 0.17]
ubtotal (95% CI) 234	4		239		♦		-0.46 [-0.64, -0.27]
est for heterogeneity: Chi² = 5.39, df = est for overall effect: Z = 4.88 (P < 0.0		, I ² = 44.4%					
04 Paroxetine							
Burnham 1993 75	9 -5.	61(7.47)	75	-4.60(7.53)			-0.13 [-0.45, 0.18]
Kamijima 2004 94	4 -8.	11(8.00)	94	-3.46(6.10)	-		-0.65 [-0.94, -0.36]
Zohar 1996a 199	8 -8.	.00(8.00)	99	-5.00(7.90)	-		-0.38 [-0.62, -0.13]
ubtotal (95% CI) 37:	1		268		•		-0.40 [-0.56, -0.23]
est for heterogeneity: Chi² = 5.56, df = est for overall effect: Z = 4.82 (P < 0.0		, I ² = 64.0%					
5 Sertraline							
Jenike 1990b	0 20.	.60(9.20)	8	22.30(7.80)		_	-0.19 [-1.12, 0.74]
Subtotal (95% CI) 10	0		8		-		-0.19 [-1.12, 0.74]
est for heterogeneity: not applicable					٦		
est for overall effect: $Z = 0.40$ (P = 0.6)	9)						
otal (95% CI) 863			766		•		-0.42 [-0.52, -0.32]
est for heterogeneity: Chi² = 13.50, df :		26), I² = 18.5%					
est for overall effect: $Z = 8.23$ (P < 0.0)	10001)						
					-4 -2 0	2 4	
					Favours SSRI	Favours placebo	





OCD adults: SSRIs Review: Comparison: 01 SSRIs vs placebo (acute phase) Outcome 18 Montgomery-Asberg Depression Rating Scale SMD (fixed) SMD (fixed) Study Placebo or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI 01 Citalopram Montgomery 2001 102 -3.40(6.30) 101 -1.30(6.70) -0.32 [-0.60, -0.04] Subtotal (95% CI) 102 101 -0.32 [-0.60, -0.04] Test for heterogeneity: not applicable Test for overall effect: Z = 2.28 (P = 0.02) Montgomery 1993 52 -2.96(8.09) 56 -2.88(9.27) -0.01 [-0.39, 0.37] Subtotal (95% CI) 52 56 -0.01 [-0.39, 0.37] Test for heterogeneity: not applicable Test for overall effect: Z = 0.05 (P = 0.96) 03 Paroxetine -0.35 [-0.60, -0.11] -0.35 [-0.60, -0.11] Zohar 1996a 198 -3.40(7.90) 99 -0.60(7.90) Subtotal (95% CI) 198 Test for heterogeneity: not applicable Test for overall effect: Z = 2.85 (P = 0.004) -0.28 [-0.44, -0.11] Test for heterogeneity: $Chi^2 = 2.42$, df = 2 (P = 0.30), $I^2 = 17.2\%$ Test for overall effect: Z = 3.30 (P = 0.0010) 0.5 Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase) Outcome 19 Hamilton Anxiety Scale SSRI SMD (fixed) SMD (fixed) Study Placebo Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Ν 01 Fluvoxamine Perse 1987 7.69(3.60) 8 12.37(7.56) -0.75 [-1.77, 0.28] Subtotal (95% CI) -0.75 [-1.77, 0.28] Test for heterogeneity: not applicable Test for overall effect: Z = 1.43 (P = 0.15) -10 10 -5 0 Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase) 20 Clinical Global Impression Outcome SSRI Placebo SMD (fixed) SMD (fixed) or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Jenike 1997 19 3.10(1.00) 18 3.20(1.00) -0.10 [-0.74, 0.55] Subtotal (95% CI) 19 18 -0.10 [-0.74, 0.55] Test for heterogeneity: not applicable Test for overall effect: Z = 0.30 (P = 0.77) -10 -5 Ó 5 10 Favours SSRI Favours placebo OCD adults: SSRIs Review: 01 SSRIs vs placebo (acute phase) Comparison: Outcome 21 Clinical Global Impressions: severity SMD (fixed) SMD (fixed) Study SSRI Placebo Mean (SD) N Mean (SD) or sub-category 01 Paroxetine -0.36 [-0.61, -0.12] -0.36 [-0.61, -0.12] Zohar 1996a -1.20(1.40) -0.70(1.30) 194 99 99 Subtotal (95% CI) 194 Test for heterogeneity: not applicable Test for overall effect: Z = 2.93 (P = 0.003) Ó Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase) 22 Sheehan Disability Scale: family Outcome Study SSRI Placebo SMD (fixed) SMD (fixed) Mean (SD) or sub-category Ν Ν Mean (SD) 95% CI 95% CI 01 Citalopram Montgomery 2001 102 -1.80(2.40) 101 -1.00(2.40) -0.33 [-0.61, -0.06] -0.33 [-0.61, -0.06] Subtotal (95% Ch. 102 101 Test for heterogeneity; not applicable Test for overall effect: Z = 2.35 (P = 0.02) -10 10 -5 Ó Favours SSRI Favours placebo

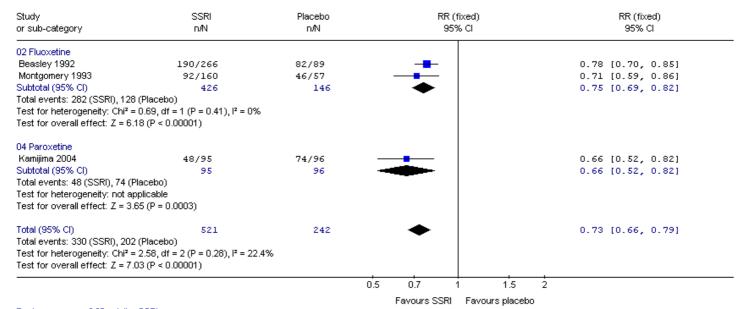
OCD adults: SSRIs Review: Comparison: 01 SSRIs vs placebo (acute phase) Outcome 23 Sheehan Disability Scale: social SMD (fixed) SMD (fixed) Study Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Citalopram -0.33 [-0.61, -0.05] -0.33 [-0.61, -0.05] Montgomery 2001 102 -2.00(2.20) 101 -1.30(2.00) Subtotal (95% CI) 102 Test for heterogeneity: not applicable Test for overall effect: Z = 2.35 (P = 0.02) -10 -5 ò 5 10 Favours SSRI Favours placebo OCD adults: SSRIs Review: 01 SSRIs vs placebo (acute phase) Comparison: Outcome 24 Sheehan Disability Scale: work SMD (fixed) SMD (fixed) Study SSRI Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Citalopram -0.35 [-0.63, -0.08] -0.35 [-0.63, -0.08] Montgomery 2001 102 -1.90(2.40)101 -1.10(2.10)Subtotal (95% CI) 101 102 Test for heterogeneity: not applicable Test for overall effect: Z = 2.50 (P = 0.01) -10 -5 Ó Favours SSRI Favours placebo Review: OCD adults: SSRIs 01 SSRIs vs placebo (acute phase) Comparison: Outcome: 25 Symptom Checklist-90 Study SSRI Placebo SMD (fixed) SMD (fixed) Ν Mean (SD) Ν or sub-category Mean (SD) 95% CI 95% CI 01 Paroxetine 169 Zohar 1996a -29.00(47.10) -19.50(37.30) -0.21 [-0.48, 0.06] Subtotal (95% CI) 169 -0.21 [-0.48, 0.06] Test for heterogeneity; not applicable Test for overall effect: Z = 1.55 (P = 0.12) -10 -5 ń 10 Favours SSRI Favours placebo OCD adults: SSRIs Review: Comparison: 01 SSRIs vs placebo (acute phase) Outcome: 26 Subanalysis: Y-BOCS in patients with comorbid depression SMD (fixed) SMD (fixed) Study SSRI Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 02 Fluoxetine Beasley 1992 75 18.90(8.30) 76 23.60(7.50) -0.59 [-0.92, -0.27] Montgomery 1993 52 -5.13(6.41) 56 -3.70(5.98) -0.23 [-0.61, 0.15] Subtotal (95% CI) 127 132 -0.44 [-0.68, -0.19] Test for heterogeneity: Chi² = 2.02, df = 1 (P = 0.16), l² = 50.4% Test for overall effect: Z = 3.47 (P = 0.0005) 03 Fluvoxamine Goodman 1989 -1.21 [-1.87, -0.54] -1.21 [-1.87, -0.54] 21 19.40(7.00) 21 28.00(7.00) Subtotal (95% CI) 21 21 Test for heterogeneity: not applicable Test for overall effect: Z = 3.56 (P = 0.0004) 04 Paroxetine Kamijima 2004 -0.65 [-0.94, -0.36] -8.11(8.00) -3.46(6.10) Subtotal (95% CI) 94 94 -0.65 [-0.94, -0.36] Test for heterogeneity: not applicable Test for overall effect: Z = 4.35 (P < 0.0001) Total (95% CI) 247 -0.58 [-0.76, -0.40] Test for heterogeneity: $Chi^2 = 6.94$, df = 3 (P = 0.07), $I^2 = 56.8\%$ Test for overall effect: Z = 6.22 (P < 0.00001)

-2

Ö Favours SSRI Favours placebo

Review: OCD adults: SSRIs Comparison: 01 SSRIs vs placebo (acute phase)

Outcome: 27 Subanalysis: Non-responders (OCD) (CGI or Y-BOCS) in patients with comorbid depression



Review: OCD adults: SSRIs

Comparison: 02 SSRIs at different doses (acute phase)

Outcome: 01 Adverse events

Study or sub-category	Dose 1 n/N	Dose 2 n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Citalopram 40mg vs Citalopra	am 20mg			
Montgomery 2001	67/98	74/102		0.94 [0.79, 1.13]
Subtotal (95% CI)	98	102		0.94 [0.79, 1.13]
otal events: 67 (Dose 1), 74 (I	Dose 2)		_	•
est for heterogeneity; not appl	licable			
est for overall effect: Z = 0.65	i (P = 0.52)			
)2 Citalopram 60mg vs Citalopra	am 20mg			
Montgomery 2001	72/100	74/102		0.99 [0.84, 1.18]
Subtotal (95% CI)	100	102		0.99 [0.84, 1.18]
otal events: 72 (Dose 1), 74 (I	Dose 2)		T	
est for heterogeneity: not appl	licable			
est for overall effect: Z = 0.09	(P = 0.93)			
3 Citalopram 60mg vs Citalopra	am 40mg			
Montgomery 2001	72/100	67/98		1.05 [0.88, 1.26]
Subtotal (95% CI)	100	98		1.05 [0.88, 1.26]
otal events: 72 (Dose 1), 67 (I	Dose 2)		-	
est for heterogeneity: not appl	licable			
est for overall effect: $Z = 0.56$	6 (P = 0.58)			
)4 Sertraline rapid titration vs s	low titration			
Bogetto 2002	13/17	11/15		1.04 [0.70, 1.56]
Subtotal (95% CI)	17	15		1.04 [0.70, 1.56]
otal events: 13 (Dose 1), 11 (I	Dose 2)			•
est for heterogeneity: not appl	licable			
est for overall effect: Z = 0.20				

Favours Dose 1 Favours Dose 2

Review: Comparison: Outcome:

OCD adults: SSRIs 02 SSRIs at different doses (acute phase) 02 Leaving study early

Study or sub-category	Dose 1 n/N	Dose 2 n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
of Citalopram 40mg vs Citalopram Monfgomery 2001 Subtotal (95% C) Total events: 15 (Dose 1), 16 (Do Fest for heterogeneity: not application of the story of the s	15/98 98 se 2) able	16/102 102	+	0.98 [0.51, 1.86] 0.98 [0.51, 1.86]
2 Citalopram 60mg vs Citalopram Montgomery 2001 Jubtotal (95% CI) otal events: 15 (Dose 1), 16 (Do est for heterogeneity: not application of the program of	15/100 100 se 2) able	16/102 102	+	0.96 [0.50, 1.83] 0.96 [0.50, 1.83]
3 Citalopram 60mg vs Citalopram Montgomery 2001 ubtotal (95% CI) otal events: 15 (Dose 1), 15 (Do est for heterogeneity: not applicates for overall effect: Z = 0.06 (F	15/100 100 se 2) able	15/98 98	+	0.98 [0.51, 1.89] 0.98 [0.51, 1.89]
4 Fluoxetine 40mg vs Fluoxetine flontgomery 1993 ubtotal (95% Cl) otal events: 13 (Dose 1), 14 (Do set for heterogeneity: not applica est for overall effect: Z = 0.17 (F	13/52 52 se 2) able	14/53 53	+	0.95 [0.49, 1.81] 0.95 [0.49, 1.81]
5 Fluoxetine 60mg vs Fluoxetine Montgomery 1993 ubtotal (95% CI) otal events: 14 (Dose 1), 14 (Do est for heterogeneity: na application of the programmer of	14/55 55 se 2) able	14/53 53	+	0.96 [0.51, 1.82] 0.96 [0.51, 1.82]
6 Fluoxetine 60mg vs Fluoxetine Montgomery 1993 Aubtotal (95% CI) otal events: 14 (Dose 1), 13 (Dose est for heterogeneity: not applicated est for overall effect: Z = 0.05 (F	40mg 14/55 55 se 2)	13/52 52	+	1.02 [0.53, 1.96] 1.02 [0.53, 1.96]
7 Paroxetine 40mg vs Paroxetine Hollander 2003d Jubtotal (95% CI) otal events: 20 (Dose 1), 14 (Dose est for heterogeneity: not applicate est for overall effect: Z = 1.21 (F	20mg 20/86 86 se 2) able	14/88 88	-	1.46 [0.79, 2.70] 1.46 [0.79, 2.70]
8 Paroxetine 60mg vs Paroxetine follander 2003d outstotal (95% Cl) otal events: 19 (Dose 1), 14 (Dose t for heterogeneity: not applicate for overall effect: Z = 1.07 (F	e 20mg 19/85 85 se 2) able	14/88 88	-	1.41 [0.75, 2.62] 1.41 [0.75, 2.62]
B Paroxetine 60mg vs Paroxetine follander 2003d ubtotal (95% Cl) stall events: 19 (Dose 1), 20 (Dose to the total events of	e 40mg 19/85 85 se 2) able	20/86 86	+	0.96 [0.55, 1.67] 0.96 [0.55, 1.67]
est for overall effect: Z = 0.14 () D Sertraline 100mg vs Sertraline Preist 1995a ubtotal (95% Cl) otal events: 27 (Dose 1), 17 (Do set for heterogeneity: not applica est for overall effect: Z = 1.69 ()	50mg 27/81 81 se 2) able	17/80 80	-	1.57 [0.93, 2.64] 1.57 [0.93, 2.64]
1 Sertraline 200mg vs Sertraline sreist 1995a ubtotal (95% CI) stal events: 21 (Dose 1), 17 (Do est for heterogeneity: not applica est for overall effect: Z = 0.74 (F	50mg 21/80 80 se 2) able	17/80 80	-	1.24 [0.71, 2.16] 1.24 [0.71, 2.16]
2 Sertraline 200mg vs Sertraline sreist 1995a ubtotal (95% CI) otal events: 21 (Dose 1), 27 (Do est for heterogeneity: not applica est for overall effect: Z = 0.88 (F	100mg 21/80 80 se 2) able	27/81 81	*	0.79 [0.49, 1.27] 0.79 [0.49, 1.27]
3 Sertraline rapid titration vs slov gogetto 2002 ubtotal (95% CI) otal events: 3 (Dose 1), 2 (Dose est for heterogeneity: not applicates est for overall effect: Z = 0.33 (F	w titration 3/17 17 2) able	2/15 15		1.32 [0.25, 6.88] - 1.32 [0.25, 6.88]

Review: OCD adults: SSRIs 02 SSRIs at different doses (acute phase) Comparison: 03 Leaving study early due to adverse events RR (fixed) RR (fixed) Dose 2 or sub-category 95% CI 95% CI 01 Citalopram 40mg vs Citalopram 20mg 4/102 1.56 [0.45, 5.36] Montgomery 2001 Subtotal (95% CI) 102 1.56 [0.45, 5.36] Total events: 6 (Dose 1), 4 (Dose 2) Test for heterogeneity; not applicable Test for overall effect: Z = 0.71 (P = 0.48) 02 Citalopram 60mg vs Citalopram 20mg Montgomery 2001 4/100 4/102 1.02 [0.26, 3.97] Subtotal (95% CI) Total events: 4 (Dose 1), 4 (Dose 2) 1.02 [0.26, 3.97] Test for heterogeneity: not applicable
Test for overall effect: Z = 0.03 (P = 0.98) 03 Citalopram 60mg vs Citalopram 40mg Montgomery 2001 Subtotal (95% CI) 0.65 [0.19, 2.24] 0.65 [0.19, 2.24] 4/100 6/98 Total events: 4 (Dose 1), 6 (Dose 2) Test for heterogeneity; not applicable Test for overall effect: Z = 0.68 (P = 0.50) 04 Fluoxetine 40mg vs Fluoxetine 20mg Beasley 1992 30/87 0.39 [0.21, 0.71] Montgomery 1993 Subtotal (95% CI) 0.51 [0.10, 2.66] 0.40 [0.23, 0.71] 2/52 140 141 Total events: 14 (Dose 1), 34 (Dose 2)
Test for heterogeneity: Chi² = 0.09, df = 1 (P = 0.77), l² = 0% Test for overall effect: Z = 3.14 (P = 0.002) 05 Fluoxetine 60mg vs Fluoxetine 20mg Beasley 1992 0.55 [0.33, 0.92] 30/87 Montgomery 1993 Subtotal (95% CI) 0.24 [0.03, 2.09] 0.51 [0.31, 0.85] 1/55 4/53 140 145 Total events: 18 (Dose 1), 34 (Dose 2) Test for heterogeneity: Chi² = 0.53, df = 1 (P = 0.46), l² = 0% Test for overall effect: Z = 2.61 (P = 0.009) 06 Fluoxetine 60mg vs Fluoxetine 40mg Beasley 1992 12/89 1.40 [0.71, 2.76] Montgomery 1993 Subtotal (95% CI) 1/55 2/52 0.47 [0.04, 5.06] 1.27 [0.66, 2.41] Total events: 18 (Dose 1), 14 (Dose 2) Test for heterogeneity: $Chi^2 = 0.75$, df = 1 (P = 0.39), $I^2 = 0\%$ Test for overall effect: Z = 0.72 (P = 0.47) 07 Paroxetine 40mg vs Paroxetine 20mg Hollander 2003d 0.91 [0.37, 2.25] Subtotal (95% CI) 88 0.91 [0.37, 2.25] Total events: 8 (Dose 1), 9 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.21 (P = 0.84) 08 Paroxetine 60mg vs Paroxetine 20mg Hollander 2003d 12/85 9/88 1.38 [0.61, 3.11] Subtotal (95% CI) Total events: 12 (Dose 1), 9 (Dose 2) 1.38 [0.61, 3.11] Test for heterogeneity: not applicable
Test for overall effect: Z = 0.78 (P = 0.44) 09 Paroxetine 60mg vs Paroxetine 40mg Hollander 2003d Subtotal (95% CI) 12/85 8/86 1.52 [0.65, 3.53] 1.52 [0.65, 3.53] Total events: 12 (Dose 1), 8 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.97 (P = 0.33) 10 Sertraline 100mg vs Sertraline 50mg Greist 1995a 11/81 6/80 1.81 [0.70, 4.66] Subtotal (95% CI) 1.81 [0.70, 4.66] Total events: 11 (Dose 1), 6 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.23 (P = 0.22) 11 Sertraline 200mg vs Sertraline 50mg Greist 1995a Subtotal (95% CI) 1.00 [0.34, 2.97] 1.00 [0.34, 2.97] 6/80 80 Total events: 6 (Dose 1), 6 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.00 (P = 1.00) 12 Sertraline 200mg vs Sertraline 100mg Greist 1995a 11/81 0.55 [0.21, 1.42] Subtotal (95% CI) 0.55 [0.21, 1.42] Total events: 6 (Dose 1), 11 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.23 (P = 0.22) 13 Sertraline rapid titration vs slow titration 2/15 Bogetto 2002 Subtotal (95% CI) 2/17 0.88 [0.14, 5.52] 0.88 [0.14, 5.52] Total events: 2 (Dose 1), 2 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.13 (P = 0.89) 0.2 0.5 1 2 5

Favours Dose 1 Favours Dose 2

OCD adults: SSRIs

Review:

02 SSRIs at different doses (acute phase) Comparison: Outcome: 04 Non-responders Dose 1 Dose 2 RR (fixed) RR (fixed) Study or sub-category n/N n/N 95% CI 95% CI 01 Citalopram 40mg vs Citalopram 20mg Montgomery 2001 47/98 44/102 1.11 [0.82, 1.51] Subtotal (95% CI) 102 1.11 [0.82, 1.51] Total events: 47 (Dose 1), 44 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.68 (P = 0.49) 02 Citalopram 60mg vs Citalopram 20mg Montgomery 2001 35/100 44/102 0.81 [0.57, 1.15] Subtotal (95% CI) 102 0.81 [0.57, 1.15] Total events: 35 (Dose 1), 44 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.18 (P = 0.24) 03 Citalopram 60mg vs Citalopram 40mg Montgomery 2001 35/100 47/98 0.73 [0.52, 1.02] Subtotal (95% CI) 100 98 0.73 [0.52, 1.02] Total events: 35 (Dose 1), 47 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.83 (P = 0.07) 04 Fluoxetine 40mg vs Fluoxetine 20mg Beasley 1992 62/87 1.02 [0.85, 1.23] 0.86 [0.66, 1.12] Montgomery 1993 33/52 39/53 Subtotal (95% CI) 140 0.96 [0.83, 1.12] 141 Total events: 98 (Dose 1), 101 (Dose 2) Test for heterogeneity: Chi² = 1.12, df = 1 (P = 0.29), l² = 10.9% Test for overall effect: Z = 0.49 (P = 0.62) 05 Fluoxetine 60mg vs Fluoxetine 20mg 0.98 [0.81, 1.19] Beasley 1992 63/90 62/87 Montgomery 1993 0.84 [0.65, 1.09] 34/55 39/53 Subtotal (95% CI) 145 140 0.93 [0.79, 1.08] Total events: 97 (Dose 1), 101 (Dose 2) Test for heterogeneity: $Chi^2 = 0.89$, df = 1 (P = 0.34), $I^2 = 0\%$ Test for overall effect: Z = 0.96 (P = 0.34) 06 Fluoxetine 60mg vs Fluoxetine 40mg Beasley 1992 63/90 65/89 0.96 [0.80, 1.15] Montgomery 1993 34/55 33/52 0.97 [0.73, 1.31] Subtotal (95% CI) 145 141 0.96 [0.82, 1.13] Total events: 97 (Dose 1), 98 (Dose 2) Test for heterogeneity: $Chi^2 = 0.01$, df = 1 (P = 0.93), $I^2 = 0\%$ Test for overall effect: Z = 0.46 (P = 0.65) 07 Paroxetine 40mg vs Paroxetine 20mg Hollander 2003d 62/88 0.84 [0.67, 1.05] Subtotal (95% CI) 88 0.84 [0.67, 1.05] Total events: 51 (Dose 1), 62 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.53 (P = 0.13) 08 Paroxetine 60mg vs Paroxetine 20mg Hollander 2003d 48/85 62/88 0.80 [0.64, 1.01] Subtotal (95% CI) 0.80 [0.64, 1.01] 85 88 Total events: 48 (Dose 1), 62 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.88 (P = 0.06) 09 Paroxetine 60mg vs Paroxetine 40mg Hollander 2003d 0.95 [0.74, 1.23] 48/85 51/86 0.95 [0.74, 1.23] Subtotal (95% CI) 85 86 Total events: 48 (Dose 1), 51 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.37 (P = 0.71) 0.7 1.5 ż 0.5 Favours Dose 1 Favours Dose 2

Review: OCD adults: SSRIs Comparison: 02 SSRIs at different doses (acute phase) Outcome: 05 Y-BOCS SMD (fixed) SMD (fixed) Study Dose 2 or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI 01 Citalopram 40mg vs Citalopram 20mg Montgomery 2001 -8.90(7.00) 102 -8.40(7.30) -0.07 [-0.35, 0.21] 98 Subtotal (95% CI) 102 -0.07 [-0.35, 0.21] Test for heterogeneity: not applicable Test for overall effect: Z = 0.49 (P = 0.62) 02 Citalopram 60mg vs Citalopram 20mg Montgomery 2001 100 -10.40(6.90) 102 -8.40(7.30) -0.28 [-0.56, 0.00] Subtotal (95% CI) -0.28 [-0.56, 0.00] 100 102 Test for heterogeneity: not applicable Test for overall effect: Z = 1.98 (P = 0.05) 03 Citalopram 60mg vs Citalopram 40mg Montgomery 2001 -0.22 [-0.49, 0.06] 100 -10.40(6.90) 98 -8.90(7.00) Subtotal (95% CI) -0.22 [-0.49, 0.06] Test for heterogeneity: not applicable Test for overall effect: Z = 1.51 (P = 0.13) 04 Fluoxetine 40mg vs Fluoxetine 20mg Beasley 1992 67 18.10(7.90) 75 18.90(8.30) -0.10 [-0.43, 0.23] Montgomery 1993 52 -4.76(6.89) 52 -5.13(6.41) 0.06 [-0.33, 0.44] Subtotal (95% CI) -0.03 [-0.28, 0.22] 119 127 Test for heterogeneity: Chi² = 0.35, df = 1 (P = 0.55), l² = 0% Test for overall effect: Z = 0.26 (P = 0.80) 05 Fluoxetine 60mg vs Fluoxetine 20mg Beasley 1992 68 16.80(7.80) 75 18.90(8.30) -0.26 [-0.59, 0.07] Montgomery 1993 54 -6.07(6.92) 52 -5.13(6.41) -0.14 [-0.52, 0.24] Subtotal (95% CI) 122 127 -0.21 [-0.46, 0.04] Test for heterogeneity: Chi² = 0.21, df = 1 (P = 0.64), l² = 0% Test for overall effect: Z = 1.63 (P = 0.10) 06 Fluoxetine 60mg vs Fluoxetine 40mg -0.16 [-0.50, 0.17] Beasley 1992 68 16.80(7.80) 67 18.10(7.90) Montgomery 1993 -6.07(6.92) 52 -0.19 [-0.57, 0.19] 54 -4.76(6.89) Subtotal (95% CI) -0.18 [-0.43, 0.08] Test for heterogeneity: Chi² = 0.01, df = 1 (P = 0.93), l² = 0% Test for overall effect: Z = 1.36 (P = 0.18) 07 Sertraline rapid titration vs slow titration Bogetto 2002 12.43(3.78) 13 13.54(4.03) -0.28 [-1.03, 0.48] Subtotal (95% CI) -0.28 [-1.03, 0.48] 14 13 Test for heterogeneity: not applicable Test for overall effect: Z = 0.71 (P = 0.48) -2 Ó Favours Dose 1 Favours Dose 2 Review: OCD adults: SSRIs Comparison: 02 SSRIs at different doses (acute phase) 06 NIMH-OC Outcome: Dose 2 SMD (fixed) SMD (fixed) Study or sub-category Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Citalopram 40mg vs Citalopram 20mg Montgomery 2001 -2.90(2.80) 102 -2.80(2.50) -0.04 [-0.31, 0.24] Subtotal (95% CI) 102 -0.04 [-0.31, 0.24] Test for heterogeneity: not applicable Test for overall effect: Z = 0.27 (P = 0.79) 02 Citalopram 60mg vs Citalopram 20mg Montgomery 2001 Subtotal (95% CI) 100 -3.10(2.60) 102 -2.80(2.50) -0.12 [-0.39, 0.16] -0.12 [-0.39, 0.16] 100 102 Test for heterogeneity: not applicable Test for overall effect: Z = 0.83 (P = 0.41) 03 Citalopram 60mg vs Citalopram 40mg Montgomery 2001 100 -3.10(2.60) -2.90(2.80) -0.07 [-0.35, 0.20] 98 Subtotal (95% CI) -0.07 [-0.35, 0.20] 98 Test for heterogeneity: not applicable Test for overall effect: Z = 0.52 (P = 0.60)

0.5

Ó Favours Dose 1 Favours Dose 2

Review: OCD adults: SSRIs

02 SSRIs at different doses (acute phase) 07 Hamilton Rating Scale for Depression Comparison: Outcome:

Study or sub-category	N	Dose 1 Mean (SD)	N	Dose 2 Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
	etine 20mg					
Montgomery 1993	52	-2.55(6.77)	52	-1.85(6.04)		-0.11 [-0.49, 0.28]
Subtotal (95% CI)	52		52			-0.11 [-0.49, 0.28]
Test for heterogeneity: not ap	oplicable					·
Test for overall effect: $Z = 0$.	55 (P = 0.58)					
02 Fluoxetine 60mg vs Fluoxe	etine 20mg					
Montgomery 1993	54	-3.41(6.23)	52	-1.85(6.04)		-0.25 [-0.63, 0.13]
Subtotal (95% CI)	54		52			-0.25 [-0.63, 0.13]
Test for heterogeneity: not ap	oplicable					
Test for overall effect: $Z = 1$.	29 (P = 0.20)					
03 Fluoxetine 60mg vs Fluoxe	etine 40mg					
Montgomery 1993	54	-3.41(6.23)	52	-2.55(6.77)		-0.13 [-0.51, 0.25]
Subtotal (95% CI)	54		52		-	-0.13 [-0.51, 0.25]
Test for heterogeneity: not ap	oplicable					
Test for overall effect: $Z = 0$.	68 (P = 0.50)					

Favours Dose 1 Favours Dose 2

Favours Dose 1 Favours Dose 2

Review:

Review: OCD adults: SSRIs
Comparison: 02 SSRIs at different doses (acute phase)
Outcome: 08 Montgomery-Asberg Depression Rating Scale

Study or sub-category	N	Dose 1 Mean (SD)	N	Dose 2 Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
	Omg					
Montgomery 2001	98	-3.90(5.20)	102	-3.40(6.30)		-0.09 [-0.36, 0.19]
Subtotal (95% CI)	98		102			-0.09 [-0.36, 0.19]
Test for heterogeneity: not applicab						
Test for overall effect: $Z = 0.61$ (P =	= 0.54)					
02 Citalopram 60mg vs Citalopram 2	Omg					
Montgomery 2001	100	-4.00(6.50)	102	-3.40(6.30)		-0.09 [-0.37, 0.18]
Subtotal (95% CI)	100		102			-0.09 [-0.37, 0.18]
Test for heterogeneity: not applicab						
Test for overall effect: Z = 0.66 (P =	0.51)					
03 Citalopram 60mg vs Citalopram 4	0mg					
Montgomery 2001	100	-4.00(6.50)	98	-3.90(5.20)	- •	-0.02 [-0.30, 0.26]
Subtotal (95% CI)	100		98			-0.02 [-0.30, 0.26]
Test for heterogeneity: not applicab						
Test for overall effect: Z = 0.12 (P =	= 0.91)					
04 Fluoxetine 40mg vs Fluoxetine 20	Omg					
Montgomery 1993	52	-5.20(8.19)	52	-2.96(8.09)		-0.27 [-0.66, 0.11]
Subtotal (95% CI)	52		52			-0.27 [-0.66, 0.11]
Test for heterogeneity: not applicab						
Test for overall effect: Z = 1.39 (P =	= 0.17)					
05 Fluoxetine 60mg vs Fluoxetine 20	Omg					
Montgomery 1993	54	-5.20(8.73)	52	-2.96(8.09)		-0.26 [-0.65, 0.12]
Subtotal (95% CI)	54		52			-0.26 [-0.65, 0.12]
Test for heterogeneity: not applicab						
Test for overall effect: Z = 1.35 (P =	0.18)					
06 Fluoxetine 60mg vs Fluoxetine 40	Omg					
Montgomery 1993	54	-5.20(8.73)	52	-5.20(8.19)		0.00 [-0.38, 0.38]
Subtotal (95% CI)	54		52			0.00 [-0.38, 0.38]
Test for heterogeneity: not applicab					T	
Test for overall effect: Z = 0.00 (P =	= 1.00)					

Review: OCD adults: SSRIs

Comparison: 02 SSRIs at different doses (acute phase)
Outcome: 09 Sheehan Disability Score: family

Study or sub-category	N	Dose 1 Mean (SD)	N	Dose 2 Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% CI
	pram 20mg					
Montgomery 2001	98	-2.10(2.30)	102	-1.80(2.40)		-0.13 [-0.40, 0.15]
Subtotal (95% CI)	98		102			-0.13 [-0.40, 0.15]
Test for heterogeneity: not a	oplicable				-	
Test for overall effect: $Z = 0$.	90 (P = 0.37)					
02 Citalopram 60mg vs Citalo	pram 20mg					
Montgomery 2001	100	-2.30(2.40)	102	-1.80(2.40)		-0.21 [-0.48, 0.07]
Subtotal (95% CI)	100		102			-0.21 [-0.48, 0.07]
est for heterogeneity: not a	oplicable				-	
est for overall effect: $Z = 1$.	47 (P = 0.14)					
03 Citalopram 60mg vs Citalo	pram 40mg					
Montgomery 2001	100	-2.30(2.40)	98	-2.10(2.30)	 _	-0.08 [-0.36, 0.19]
Subtotal (95% CI)	100		98			-0.08 [-0.36, 0.19]
Fest for heterogeneity: not a	oplicable				7	
Fest for overall effect: $Z = 0$.	60 (P = 0.55)					

Favours Dose 1 Favours Dose 2

Favours Dose 1 Favours Dose 2

Favours Dose 1 Favours Dose 2

Review: OCD adults: SSRIs

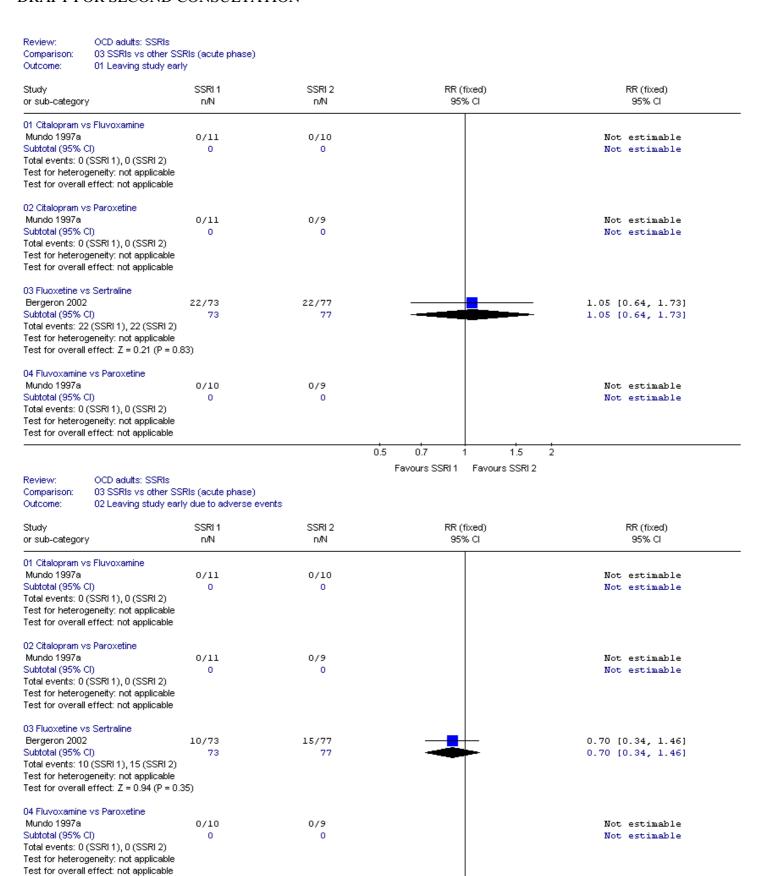
Comparison: 02 SSRIs at different doses (acute phase)
Outcome: 10 Sheehan Disability Score: social

Study or sub-category	N	Dose 1 Mean (SD)	N	Dose 2 Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% CI
01 Citalopram 40mg vs Citalop	oram 20mg					
Montgomery 2001	98	-1.90(2.30)	102	-2.00(2.20)	 _	0.04 [-0.23, 0.32]
Subtotal (95% CI)	98		102			0.04 [-0.23, 0.32]
Fest for heterogeneity: not ap	plicable				Γ	
Test for overall effect: $Z = 0.3$						
02 Citalopram 60mg vs Citalop	oram 20mg					
Montgomery 2001	100	-2.40(2.50)	102	-2.00(2.20)		-0.17 [-0.45, 0.11]
Subtotal (95% CI)	100		102			-0.17 [-0.45, 0.11]
est for heterogeneity; not ap	plicable				-	
est for overall effect: $Z = 1.5$	20 (P = 0.23)					
03 Citalopram 60mg vs Citalop	oram 40mg					
Montgomery 2001	100	-2.40(2.50)	98	-1.90(2.30)		-0.21 [-0.49, 0.07]
Subtotal (95% CI)	100		98		-	-0.21 [-0.49, 0.07]
Fest for heterogeneity: not ap	plicable					
Test for overall effect: Z = 1.4	45 (P = 0.15)					

teview: OCD adults: SSRIs

Comparison: 02 SSRIs at different doses (acute phase)
Outcome: 11 Sheehan Disability Score: work

Study Dose 2 SMD (fixed) SMD (fixed) or sub-category Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Citalopram 40mg vs Citalopram 20mg Montgomery 2001 -2.10(2.30) 102 -1.90(2.40) -0.08 [-0.36, 0.19] Subtotal (95% CI) 102 -0.08 [-0.36, 0.19] Test for heterogeneity: not applicable Test for overall effect: Z = 0.60 (P = 0.55) 02 Citalopram 60mg vs Citalopram 20mg 100 -0.04 [-0.32, 0.24] -0.04 [-0.32, 0.24] Montgomery 2001 Subtotal (95% CI) -2.00(2.50) 102 -1.90(2.40)100 102 Test for heterogeneity: not applicable Test for overall effect: Z = 0.29 (P = 0.77) 03 Citalopram 60mg vs Citalopram 40mg Montgomery 2001 0.04 [-0.24, 0.32] 100 -2.00(2.50) 98 -2.10(2.30) Subtotal (95% CI) 100 98 0.04 [-0.24, 0.32] Test for heterogeneity: not applicable Test for overall effect: Z = 0.29 (P = 0.77) 0.5



0.2

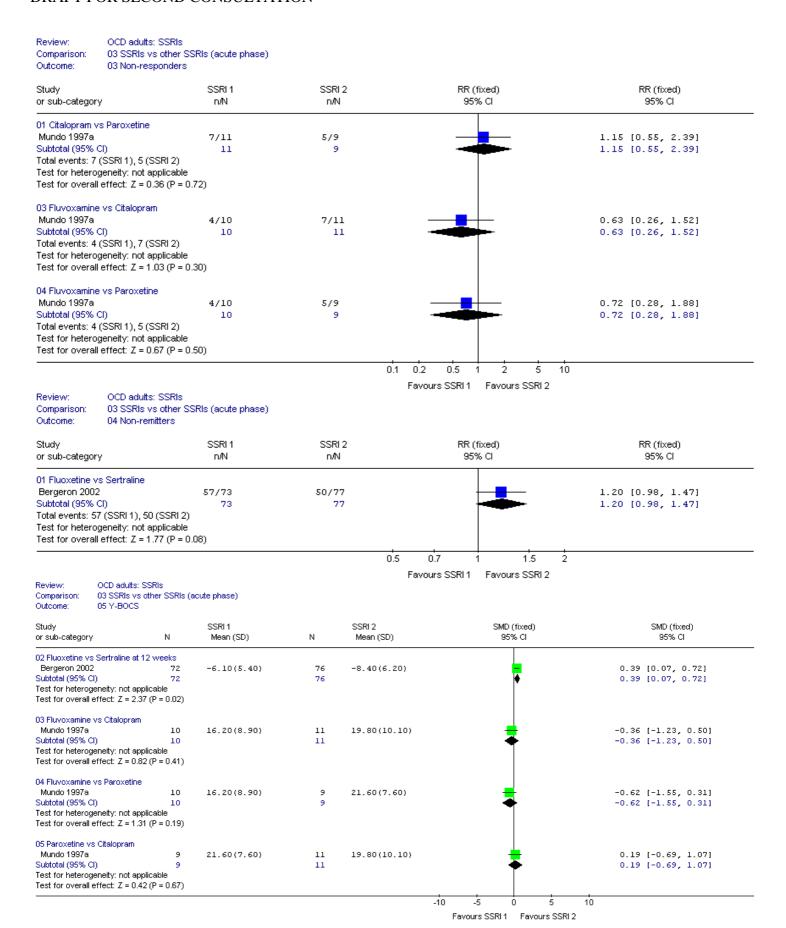
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Favours SSRI1

ż

Favours SSRI 2

5 10

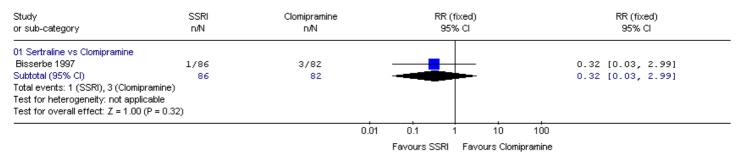


Review: OCD adults: SSRIs Comparison: 03 SSRIs vs other SSRIs (acute phase) Outcome 06 NIMH-OC SMD (fixed) SSRI 2 SMD (fixed) Study SSRI 1 or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI 01 Fluoxetine vs Sertraline Bergeron 2002 72 5.70(2.60) 76 5.50(2.80) 0.07 [-0.25, 0.40] Subtotal (95% CI) 72 0.07 [-0.25, 0.40] Test for heterogeneity: not applicable Test for overall effect: Z = 0.45 (P = 0.65) 02 Fluvoxamine vs Citalopram Mundo 1997a 10 6.23(1.81) 11 8.58(1.88) -1.22 [-2.17, -0.27] Subtotal (95% CI) 10 11 -1.22 [-2.17, -0.27] Test for heterogeneity; not applicable Test for overall effect: Z = 2.52 (P = 0.01) 03 Fluvoxamine vs Paroxetine Mundo 1997a 10 6.23(1.81) 9 7.77(2.02) -0.77 [-1.71, 0.17] Subtotal (95% CI) -0.77 [-1.71, 0.17] 10 Test for heterogeneity: not applicable Test for overall effect: Z = 1.60 (P = 0.11)04 Paroxetine vs Citalopram Mundo 1997a 9 7.77(2.02) 8.58(1.88) -0.40 [-1.29, 0.49] 11 Subtotal (95% CI) 1.1 -0.40 [-1.29, 0.49] Test for heterogeneity: not applicable Test for overall effect: Z = 0.88 (P = 0.38) Favours SSRI1 Favours SSRI2 Review: OCD adults: SSRIs Comparison: 03 SSRIs vs other SSRIs (acute phase) Outcome 07 Hamilton Rating Scale for Depression SMD (fixed) Study SSRI1 SSRI 2 SMD (fixed) or sub-category Ν Mean (SD) M Mean (SD) 95% CI 95% CI 01 Fluoxetine vs Sertraline Bergeron 2002 7.10(5.40) 6.20(4.60) 0.18 [-0.14, 0.50] 72 76 Subtotal (95% CI) 76 0.18 [-0.14, 0.50] 72 Test for heterogeneity: not applicable Test for overall effect: Z = 1.09 (P = 0.28)-0.5 0.5 Ó Favours SSRI1 Favours SSRI2 Review: OCD adults: SSRIs Comparison: 04 SSRIs vs Clomipramine (acute phase) Outcome: 01 Adverse events SSRI RR (fixed) RR (fixed) Study Clomipramine 95% CI 95% CI or sub-category n/Ν n/N 01 Fluvoxamine vs Clomipramine Freeman 1994 30/34 24/30 1.10 [0.89, 1.37] Koran 1996a 35/37 39/41 0.99 [0.90, 1.10] 0.94 [0.80, 1.10] Mundo 2001 81/115 84/112 Subtotal (95% CI) 186 183 0.98 [0.89, 1.09] Total events: 146 (SSRI), 147 (Clomipramine) Test for heterogeneity: Chi² = 1.47, df = 2 (P = 0.48), l² = 0% Test for overall effect: Z = 0.36 (P = 0.72) 02 Paroxetine vs Clomipramine Burnham 1993 77/82 80/82 0.96 [0.90, 1.03] 0.94 [0.85, 1.05] Zohar 1996a 163/201 85/99 Subtotal (95% CI) 283 181 0.95 [0.89, 1.02] Total events: 240 (SSRI), 165 (Clomipramine) Test for heterogeneity: $Chi^2 = 0.13$, df = 1 (P = 0.72), $I^2 = 0\%$ Test for overall effect: Z = 1.45 (P = 0.15) 03 Sertraline vs Clomipramine Askin 1999 7/18 13/18 0.54 [0.28, 1.03] Bisserbe 1997 56/86 57/82 0.94 [0.76, 1.16] Subtotal (95% CI) 100 0.86 [0.71, 1.06] 104 Total events: 63 (SSRI), 70 (Clomipramine) Test for heterogeneity: Chi² = 2.62, df = 1 (P = 0.11), l² = 61.8% Test for overall effect: Z = 1.41 (P = 0.16) Total (95% CI) 0.95 [0.89, 1.00] 464 Total events: 449 (SSRI), 382 (Clomipramine) Test for heterogeneity: $Chi^2 = 5.93$, df = 6 (P = 0.43), $I^2 = 0\%$ Test for overall effect: Z = 1.80 (P = 0.07)0.5 0.7 1.5 Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)

Outcome: 02 Attempted suicide



Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)

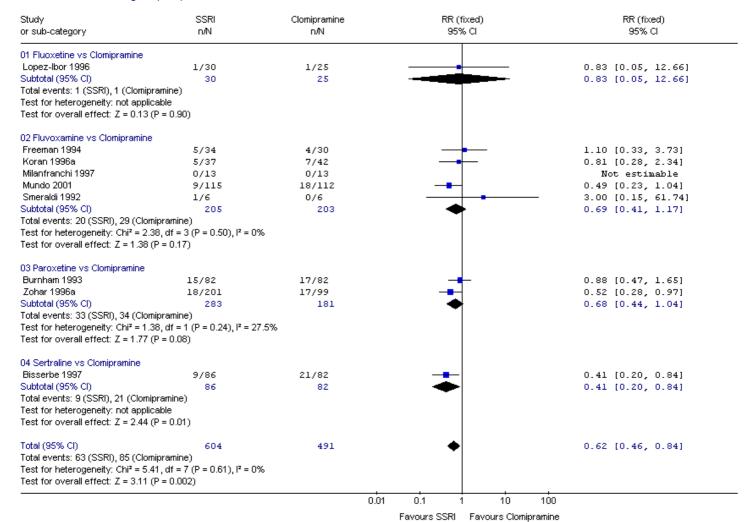
Outcome: 03 Leaving study early

Study	SSRI	Clomipramine	RR (fixed)	RR (fixed)
or sub-category	n/N	n/N	95% CI	95% CI
01 Fluoxetine vs Clomipramine				
Lopez-lbor 1996	5/30	4/25		1.04 [0.31, 3.47]
Subtotal (95% CI)	5/30	4/25		1.04 [0.31, 3.47]
Test for heterogeneity: not ap	plicable		_ _	,
Test for overall effect: $Z = 0.0$	07 (P = 0.95)			
02 Fluvoxamine vs Clomipram	ine			
Freeman 1994	6/34	11/32		0.51 [0.22, 1.23]
Koran 1996a	8/37	15/42		0.61 [0.29, 1.26]
Milanfranchi 1997	0/13	1/13	•	- 0.33 [0.01, 7.50]
Mundo 2001	19/115	26/112		0.71 [0.42, 1.21]
Smeraldi 1992	1/6	1/6		1.00 [0.08, 12.56]
Subtotal (95% CI)	34/205	54/205	-	0.64 [0.44, 0.93]
Test for heterogeneity: Chi ² =	•	•	•	,,
Test for overall effect: Z = 2.3				
03 Paroxetine vs Clomipramin	e			
Burnham 1993	28/82	28/82	_	1.00 [0.65, 1.53]
Zohar 1996a	49/201	34/99		0.71 [0.49, 1.02]
Subtotal (95% CI)	77/283	62/181	-	0.82 [0.62, 1.08]
Test for heterogeneity: Chi ² =	•		<u> </u>	,
Test for overall effect: Z = 1.4				
04 Sertraline vs Clomipramine				
Askin 1999	2/20	4/22 -	<u> </u>	0.55 [0.11, 2.69]
Bisserbe 1997	23/86	35/82		0.63 [0.41, 0.96]
Subtotal (95% CI)	25/106	39/104	-	0.62 [0.41, 0.94]
Test for heterogeneity: Chi² =	-		-	
Test for overall effect: Z = 2.2				
Total (95% CI)	141/624	159/515	•	0.72 [0.59, 0.88]
Test for heterogeneity: Chi ² =	4.26, $df = 9 (P = 0.89), I^2 = 0$	%	7	,
Test for overall effect: Z = 3.2				
	•	0.1	0.2 0.5 1 2 5	10

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)
Outcome: 04 Leaving study early due to adverse events



Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clonipramine (acute phase)
Outcome: 05 Non-responders

Study	SSRI	Clomipramine	RR (fixed)	RR (fixed)
r sub-category	n/N	n/N	95% Cl	95% CI
1 Fluoxetine vs Clomipramine				
Lopez-lbor 1996	22/30	11/25	-	1.67 [1.02, 2.73]
Subtotal (95% CI)	30	25	-	1.67 [1.02, 2.73]
otal events: 22 (SSRI), 11 (Clom				
est for heterogeneity: not applic	cable			
est for overall effect: Z = 2.03 ((P = 0.04)			
2 Fluvoxamine vs Clomipramine				
(oran 1996a	18/37	21/42		0.97 [0.62, 1.52]
Milanfranchi 1997	2/13	3/13 -		0.67 [0.13, 3.35]
Mundo 2001	46/115	44/112		1.02 [0.74, 1.40]
ubtotal (95% CI)	165	167	•	0.99 [0.76, 1.28]
otal events: 66 (SSRI), 68 (Clori			T	,
est for heterogeneity: Chi ² = 0.2	, ,	0%		
est for overall effect: Z = 0.08 (
3 Paroxetine vs Clomipramine				
Burnham 1993	47/82	42/82	- 	1.12 [0.84, 1.48]
Zohar 1996a	92/201	47/99	-	0.96 [0.75, 1.25]
ubtotal (95% CI)	283	181	•	1.03 [0.85, 1.24]
otal events: 139 (SSRI), 89 (Clo	mipramine)		ſ	
est for heterogeneity: $Chi^2 = 0.5$ est for overall effect: $Z = 0.27$ (, , , , ,	0%		
est for overall effect. Z = 0.27 (,r = 0.79)			
4 Sertraline ∨s Clomipramine ∃isserbe 1997	24 (05	20 (02	_	0.70 (0.50 1.24)
ubtotal (95% CI)	24/86 86	29/82		0.79 [0.50, 1.24] 0.79 [0.50, 1.24]
		82		0.79 [0.80, 1.24]
otal events: 24 (SSRI), 29 (Clom est for heterogeneity: not applic				
- ,				
est for overall effect: Z = 1.04 ((P = 0.30)			
otal (95% CI)	564	455	•	1.02 [0.89, 1.17]
otal events: 251 (SSRI), 197 (CI	lomipramine)		ſ	• •
est for heterogeneity: Chi² = 6.0	01 , df = 6 (P = 0.42), l ² = 0	0.1%		
est for overall effect: Z = 0.24 (
		0.1	0.2 0.5 1 2 5	10
			Favours SSRI Favours Clomip	ramine
eview: OCD adults: SSRIs				

Review: OCD adults: SSRIs
Comparison: 04 SSRIs vs Clomipramine (acute phase)
Outcome: 06 Y-BOCS

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)) (fixed) 5% Cl	SMD (fixed) 95% Cl
01 Fluoxetine vs Clomipramine							
Lopez-Ibor 1996	30	-7.50(9.29)	24	-8.90(7.13)		-	0.16 [-0.37, 0.70]
Subtotal (95% CI)	30		24				0.16 [-0.37, 0.70]
Test for heterogeneity: not app	licable						
Test for overall effect: Z = 0.60) (P = 0.55)						
02 Fluvoxamine vs Clomipramin	ie						
Koran 1996a	34	17.80(7.70)	39	17.00(8.55)		 • 	0.10 [-0.36, 0.56]
Milanfranchi 1997	13	18.40(9.20)	12	16.50(11.00)		 •	0.18 [-0.60, 0.97]
Mundo 2001	67	14.70(10.10)	61	12.50(8.00)	-	 	0.24 [-0.11, 0.59]
Smeraldi 1992	5	18.20(2.50)	5	15.20(12.40)		<u> </u>	0.30 [-0.95, 1.55]
Subtotal (95% CI)	119		117				0.19 [-0.06, 0.45]
Test for heterogeneity: Chi² = 0	1.26, df = 3 (P	= 0.97), I ² = 0%					
Test for overall effect: $Z = 1.46$	S (P = 0.14)						
03 Paroxetine vs Clomipramine							
Burnham 1993	79	-5.61(7.47)	78	-7.73(7.42)		-	0.28 [-0.03, 0.60]
Zohar 1996a	198	-8.00(8.00)	94	-8.00(8.20)	_	+	0.00 [-0.25, 0.25]
Subtotal (95% CI)	277		172			-	0.11 [-0.09, 0.30]
Test for heterogeneity: Chi² = 1	.94, df = 1 (P	= 0.16), I ² = 48.4%					
Test for overall effect: $Z = 1.09$	9 (P = 0.28)						
Total (95% CI)	426		313			•	0.14 [-0.01, 0.29]
Test for heterogeneity: Chi² = 2		= 0.87), I ² = 0%					
Test for overall effect: $Z = 1.85$	(P = 0.06)						
					1 -0.5	0 0.5 1	
					Favours SSRI	Favours Clomipramine	

Review: OCD adults: SSRIs

04 SSRIs vs Clomipramine (acute phase) 07 NIMH-OC Comparison:

Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)		SMD (95%		SMD (fixed) 95% Cl	
01 Fluvoxamine vs Clomipro	amine								
Mundo 2001	112	5.90(3.10)	105	5.90(2.80)				0.00 [-0.27, 0.2	27]
Subtotal (95% CI)	112		105					0.00 [-0.27, 0.2	27]
Test for heterogeneity: not	applicable					1	-	·	
Test for overall effect: Z =									
02 Paroxetine vs Clomipran	nine								
Burnham 1993	79	-1.42(2.04)	78	-2.06(2.03)			-	0.31 [0.00, 0.63	3]
Zohar 1996a	198	-2.50(2.90)	94	-2.50(2.60)				0.00 [-0.25, 0.2	25]
Subtotal (95% CI)	277		172			-	•	0.12 [-0.08, 0.3	31]
Test for heterogeneity: Chi-	² = 2.36, df = 1 (P	= 0.12), I ² = 57.6%					-	•	
Test for overall effect: Z =	1.20 (P = 0.23)								
Total (95% CI)	389		277			•	•	0.08 [-0.08, 0.2	23]
Test for heterogeneity: Chi-	2 = 2.86, df = 2 (P	= 0.24), I ² = 30.0%					-		
Test for overall effect: Z =	0.97 (P = 0.33)								
					-1	-0.5	0.5	1	
					F	avours SSRI	Favours Clor	mipramine	

Review:

Review: OCD adults: SSRIs
Comparison: 04 SSRIs vs Clomipramine (acute phase)
Outcome: 08 Comprehensive Psychopathological Rating Scale: obsessive-compulsive

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)		SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Fluoxetine vs Clomipran	nine						
Lopez-lbor 1996	30	-8.10(7.64)	24	-10.90(7.33)			O.37 [-0.17, 0.91]
Subtotal (95% CI)	30		24				0.37 [-0.17, 0.91]
Test for heterogeneity: not	applicable						
Test for overall effect: Z =	1.33 (P = 0.18)						
					-1	-0.5 0 0.5	+

Favours SSRI Favours Clomipramine Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase) 09 Hamilton Rating Scale for Depression Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl	
01 Fluoxetine vs Clomipramin	е						
Lopez-lbor 1996	30	-7.50(5.93)	24	-9.40(7.71)		0.28 [-0.26, 0.82]	
Subtotal (95% CI)	30		24			0.28 [-0.26, 0.82]	
Test for heterogeneity: not ap	plicable						
Test for overall effect: $Z = 1$.	00 (P = 0.32)						
02 Fluvoxamine vs Clomipram	ine						
Koran 1996a	34	6.10(3.30)	39	5.80(4.70)		0.07 [-0.39, 0.53]	
Mundo 2001	112	7.70(6.00)	105	7.00(4.50)	- 	0.13 [-0.14, 0.40]	
Subtotal (95% CI)	146		144			0.12 [-0.11, 0.35]	
Test for heterogeneity: Chi² =	0.05, df = 1 (P	= 0.83), I ² = 0%			-		
Test for overall effect: $Z = 0$.	99 (P = 0.32)						
Total (95% CI)	176		168			0.14 [-0.07, 0.35]	
Test for heterogeneity: Chi² =	0.33, df = 2 (P	= 0.85), I ² = 0%			-		
Test for overall effect: $Z = 1$:	30 (P = 0.19)						

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs Comparison: 04 SSRIs vs Clomipramine (acute phase) Outcome 10 Montgomery-Asberg Depression Rating Scale SMD (fixed) SMD (fixed) Study Clomipramine or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine vs Clomipramine Lopez-lbor 1996 30 -10.10(8.20) 24 -13.80(9.09) 0.42 [-0.12, 0.97] Subtotal (95% CI) 30 24 0.42 [-0.12, 0.97] Test for heterogeneity: not applicable Test for overall effect: Z = 1.53 (P = 0.13) 02 Fluvoxamine vs Clomipramine Smeraldi 1992 5 11.80(6.80) 5 9.60(10.30) 0.23 [-1.02, 1.47] Subtotal (95% CI) 0.23 [-1.02, 1.47] 5 Test for heterogeneity: not applicable Test for overall effect: Z = 0.36 (P = 0.72) 03 Paroxetine vs Clomipramine Zohar 1996a 198 -3.40(7.90) 94 -2.40(7.30) -0.13 [-0.38, 0.12] Subtotal (95% CI) -0.13 [-0.38, 0.12] 198 Test for heterogeneity: not applicable Test for overall effect: Z = 1.03 (P = 0.30) 123 -0.03 [-0.25, 0.19] Test for heterogeneity: $Chi^2 = 3.47$, df = 2 (P = 0.18), $I^2 = 42.4\%$ Test for overall effect: Z = 0.24 (P = 0.81) Favours SSRI Favours Clomipramine Review: OCD adults: SSRIs Comparison: 04 SSRIs vs Clomipramine (acute phase) Outcome 11 Depression (HRSD; MADRS) SMD (fixed) SMD (fixed) Study SSRI Clomipramine Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Ν 01 Fluvoxamine vs Clomipramine Koran 1996a 34 6.10(3.30) 39 5.80(4.70) 0.07 [-0.39, 0.53] Mundo 2001 0.13 [-0.14, 0.40] 112 7.70(6.00) 105 7.00(4.50) Smeraldi 1992 5 11.80(6.80) 9.60(10.30) 0.23 [-1.02, 1.47] Subtotal (95% CI) 151 149 0.12 [-0.11, 0.35] Test for heterogeneity: $Chi^2 = 0.08$, df = 2 (P = 0.96), $I^2 = 0\%$ Test for overall effect: Z = 1.04 (P = 0.30)02 Paroxetine vs Clomipramine Zohar 1996a 198 -3.40(7.90) 94 -2.40(7.30)-0.13 [-0.38, 0.12] Subtotal (95% CI) -0.13 [-0.38, 0.12] 198 94 Test for heterogeneity: not applicable Test for overall effect: Z = 1.03 (P = 0.30) Total (95% CI) 243 0.01 [-0.16, 0.17] Test for heterogeneity: Chi² = 2.21, df = 3 (P = 0.53), l² = 0% Test for overall effect: Z = 0.06 (P = 0.95) -4 -2 Ò Favours SSRI Favours Clomipramine OCD adults: SSRIs Review: Comparison: 04 SSRIs vs Clomipramine (acute phase) 12 Clinical Anxiety Scale Outcome SMD (fixed) SMD (fixed) or sub-category M Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluvoxamine vs Clomipramine Mundo 2001 0.07 [-0.20, 0.33] 7.00(4.60) 6.70(4.30) Subtotal (95% CI) 105 0.07 [-0.20, 0.33] 112 Test for heterogeneity: not applicable Test for overall effect: Z = 0.49 (P = 0.62) -0.5 Ó 0.5 Favours SSRI Favours Clomipramine Review: OCD adults: SSRIs 04 SSRIs vs Clomipramine (acute phase) Comparison: Outcome 13 Covi Anxiety Scale Study SMD (fixed) SMD (fixed) SSRI Clomipramine or sub-category Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine vs Clomipramine 0.00 [-0.54, 0.54] Lopez-lbor 1996 30 -2.40(3.10) 24 -2.40(3.09) Subtotal (95% CI) 0.00 [-0.54, 0.54] 30 24 Test for heterogeneity: not applicable

-0.5

Favours SSRI

Ó

0.5

Favours Clomipramine

Test for overall effect: Z = 0.00 (P = 1.00)

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)

Outcome: 14 Anxiety (Clinical Anxiety Scale; Covi Anxiety Scale)

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Fluoxetine vs Clomipramir	ne					
Lopez-lbor 1996	30	-2.40(3.10)	24	-2.40(3.09)		0.00 [-0.54, 0.54]
Subtotal (95% CI)	30		24			0.00 [-0.54, 0.54]
Test for heterogeneity: not a	applicable				T	
Test for overall effect: Z = 0	0.00 (P = 1.00)					
02 Fluvoxamine vs Clomiprai	mine					
Mundo 2001	112	7.00(4.60)	105	6.70(4.30)		0.07 [-0.20, 0.33]
Subtotal (95% CI)	112		105			0.07 [-0.20, 0.33]
Fest for heterogeneity: not a	applicable				_	
Test for overall effect: Z = 0).49 (P = 0.62)					
Total (95% CI)	142		129			0.05 [-0.18, 0.29]
Test for heterogeneity: Chi ²	= 0.05, df = 1 (P	= 0.83), I ² = 0%				
Test for overall effect: $Z = 0$	1.44 (P = 0.66)					

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase) 15 Clinical Global Impressions: global improvement Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
	e					
Lopez-lbor 1996	30	-2.60(1.13)	24	-2.60(1.19)		0.00 [-0.54, 0.54]
Subtotal (95% CI)	30		24			0.00 [-0.54, 0.54]
Test for heterogeneity: not a	pplicable				T	
Test for overall effect: $Z = 0$	00 (P = 1.00)					
02 Fluvoxamine vs Clomipran	nine					
Mundo 2001	112	2.30(1.00)	105	2.30(1.00)		0.00 [-0.27, 0.27]
Subtotal (95% CI)	112		105			0.00 [-0.27, 0.27]
Test for heterogeneity: not a	pplicable				Ī	
Test for overall effect: $Z = 0$.	00 (P = 1.00)					
Total (95% CI)	142		129		•	0.00 [-0.24, 0.24]
Test for heterogeneity: Chi ² :	0.00, df = 1 (P	= 1.00), I ² = 0%			T	
Test for overall effect: Z = 0	00 (P = 1.00)					
					-0.5 0 0.5	<u>+</u>

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

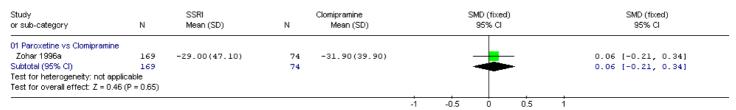
04 SSRIs vs Clomipramine (acute phase) 16 Clinical Global Impressions: severity of illness Comparison: Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)			D (fixed) 15% Cl			SMD (fixed) 95% Cl
01 Fluoxetine vs Clomipramin	e									
Lopez-lbor 1996	30	-0.90(1.23)	24	-1.90(1.14)				-		0.83 [0.27, 1.39]
Subtotal (95% CI)	30		24				•	•		0.83 [0.27, 1.39]
Test for heterogeneity: not ap	pplicable						-			
Test for overall effect: $Z = 2$.	89 (P = 0.004)									
02 Fluvoxamine vs Clomipram	nine									
Mundo 2001	112	3.40(1.30)	105	3.20(1.20)			 -			0.16 [-0.11, 0.43]
Subtotal (95% CI)	112		105				♦			0.16 [-0.11, 0.43]
Test for heterogeneity: not ap	pplicable									
Test for overall effect: $Z = 1$.	17 (P = 0.24)									
03 Paroxetine vs Clomipramin	ne									
Zohar 1996a	194	-1.20(1.40)	94	-1.10(1.20)			#			-0.07 [-0.32, 0.17]
Subtotal (95% CI)	194		94				•			-0.07 [-0.32, 0.17]
Test for heterogeneity: not ap	pplicable									
Test for overall effect: $Z = 0$.	59 (P = 0.55)									
Total (95% CI)	336		223				•			0.11 [-0.06, 0.28]
Test for heterogeneity: Chi² =	8.57, df = 2 (P	= 0.01), I ² = 76.7%					[
Test for overall effect: $Z = 1$.	23 (P = 0.22)									
					-4	-2	ó	2	4	
					Fa	vours SSR	l Favo	urs Clomi	ramine	

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)

17 Symptom Checklist-90 Outcome:



Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 04 SSRIs vs Clomipramine (acute phase)

Outcome: 18 Subanalysis: Non-responders in patients with comorbid depression

Study or sub-category	SSRI n/N	Clomipramine n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Fluoxetine vs Clomipramine				
Lopez-lbor 1996	22/30	11/25	─ ■	1.67 [1.02, 2.73]
Subtotal (95% CI)	30	25	-	1.67 [1.02, 2.73]
Total events: 22 (SSRI), 11 (Clomi	pramine)		-	
Test for heterogeneity; not applica	able			
Test for overall effect: Z = 2.03 (F	9 = 0.04)			
02 Fluvoxamine vs Clomipramine				
Mundo 2001	46/115	44/112	-	1.02 [0.74, 1.40]
Subtotal (95% CI)	115	112	•	1.02 [0.74, 1.40]
Total events: 46 (SSRI), 44 (Clomi	pramine)		T	
Test for heterogeneity; not applica	able			
Test for overall effect: Z = 0.11 (F	9 = 0.91)			
Total (95% CI)	145	137	•	1.16 [0.88, 1.51]
Total events: 68 (SSRI), 55 (Clomi	pramine)		*	. , ,
Test for heterogeneity: Chi ² = 2.72	•	3.3%		
Test for overall effect: Z = 1.06 (F				
		0.1	0.2 0.5 1 2 5	10

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison:

04 SSRIs vs Clomipramine (acute phase)
19 Subanalysis: Y-BOCS in patients with comorbid depression Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)			D (fixed) 95% Cl			SMD (fixed) 95% Cl
01 Fluoxetine vs Clomiprami	ine									
Lopez-lbor 1996	30	-7.50(9.29)	24	-8.90(7.13)			-			0.16 [-0.37, 0.70]
Subtotal (95% CI)	30		24					_		0.16 [-0.37, 0.70]
Test for heterogeneity: not a	applicable									
Test for overall effect: $Z = 0$	0.60 (P = 0.55)									
02 Fluvoxamine vs Clomipra	amine									
Mundo 2001	67	14.70(10.10)	61	12.50(8.00)			+-			0.24 [-0.11, 0.59]
Smeraldi 1992	5	18.20(2.50)	5	15.20(12.40)			-		→	0.30 [-0.95, 1.55]
Subtotal (95% CI)	72		66				-			0.24 [-0.09, 0.58]
Test for heterogeneity: Chi²	= 0.01, df $= 1$ (P	= 0.92), I ² = 0%					-			
Test for overall effect: $Z = 1$	1.42 (P = 0.16)									
Total (95% CI)	102		90					-		0.22 [-0.06, 0.51]
Test for heterogeneity: Chi²	= 0.07, df $= 2$ (P	= 0.97), I ² = 0%					_			
Test for overall effect: $Z = 1$	1.52 (P = 0.13)									
					-1	-0.5	ó	0.5	i	
						Favours SSF	RI Favour	rs Clomi;	oramine	

Review: OCD adults: SSRIs

Comparison:

04 SRIs vs Clonipramine (acute phase)
20 Subanalysis: Hamilton Rating Scale for Depression in patients with comorbid depression Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)		SMD (fix 95% C	,		SMD (fixed) 95% Cl
01 Fluoxetine vs Clomiprai	mine								
Lopez-lbor 1996	30	-7.50(5.93)	24	-9.40(7.71)			-		0.28 [-0.26, 0.82]
Subtotal (95% CI)	30		24						0.28 [-0.26, 0.82]
Test for heterogeneity: no	t applicable						_		
Test for overall effect: Z =	= 1.00 (P = 0.32)								
02 Fluvoxamine vs Clomip	ramine								
Mundo 2001	112	7.70(6.00)	105	7.00(4.50)		-			0.13 [-0.14, 0.40]
Subtotal (95% CI)	112		105			-			0.13 [-0.14, 0.40]
Test for heterogeneity: no	t applicable					-			
Test for overall effect: Z =	= 0.96 (P = 0.34)								
Total (95% CI)	142		129						0.16 [-0.08, 0.40]
Test for heterogeneity: Ch	$ni^2 = 0.22$, $df = 1$ (P	= 0.64), I ² = 0%				-			,
Test for overall effect: Z =	= 1.31 (P = 0.19)								
					-1	-0.5 0	0.5	1	
					Fa	vours SSRI F	avours Clomip	ramine	

OCD adults: SSRIs

05 SSRIs/Clomipramine vs Non-SRIs (acute phase) 01 Y-BOCS Comparison:

Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Other drug Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Fluoxetine vs Phenelzine						
Jenike 1997	19	16.20(6.30)	17	16.30(7.00)		-0.01 [-0.67, 0.64]
Subtotal (95% CI)	19		17		•	-0.01 [-0.67, 0.64]
Test for heterogeneity: not appl	icable					
Test for overall effect: $Z = 0.04$	(P = 0.96)					
02 Fluvoxamine vs Desipramine	:					
Goodman 1990a	21	19.50(9.00)	19	25.80(6.00)		-0.80 [-1.45, -0.15]
Subtotal (95% CI)	21		19		•	-0.80 [-1.45, -0.15]
Test for heterogeneity: not appl						
Test for overall effect: Z = 2.42	(P = 0.02)					
03 Clomipramine vs Buspirone						
Pato 1991	9	17.67(4.97)	9	16.78(6.46)	- -	0.15 [-0.78, 1.07]
Subtotal (95% CI)	9		9			0.15 [-0.78, 1.07]
Test for heterogeneity: not appl	icable					
Test for overall effect: $Z = 0.31$	(P = 0.76)					
04 Sertraline vs Desipramine						
Hoehn-Saric 2000	79	-8.40(8.00)	85	-6.00(8.30)		-0.29 [-0.60, 0.02]
Subtotal (95% Cl)	79		85		◆	-0.29 [-0.60, 0.02]
Test for heterogeneity: not appl	icable					
Test for overall effect: $Z = 1.86$	(P = 0.06)					
Total (95% CI)	128		130		•	-0.30 [-0.54, -0.05]
Test for heterogeneity: Chi² = 3	.91, df = 3 (P	= 0.27), I ² = 23.3%			-	
Test for overall effect: Z = 2.35	(P = 0.02)					

Favours SSRI Favours other drug

OCD adults: SSRIs Review:

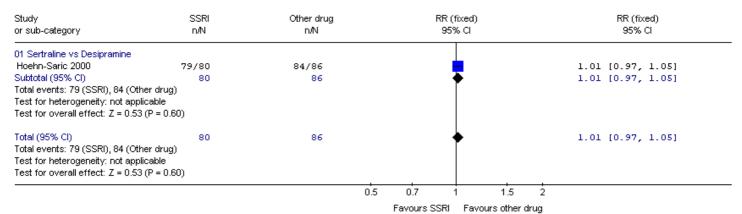
Comparison: Outcome: 05 SSRIs/Clomipramine vs Non-SRIs (acute phase) 02 NIMH-OC

Study or sub-category	N	SSRI Mean (SD)	N	Other drug Mean (SD)		SMD (fixed) 95% Cl	SMD (fixed) 95% CI
01 Fluoxetine vs Phenelzine							
Jenike 1997	19	7.00(1.10)	17	6.80(1.40)			0.16 [-0.50, 0.81]
Subtotal (95% CI)	19		17				0.16 [-0.50, 0.81]
Test for heterogeneity: not app	olicable						
est for overall effect: Z = 0.4	7 (P = 0.64)						
02 Sertraline vs Desipramine							
Hoehn-Saric 2000	79	-2.70(2.67)	85	-1.70(2.77)			-0.37 [-0.67, -0.06]
Subtotal (95% CI)	79		85				-0.37 [-0.67, -0.06]
Fest for heterogeneity: not app	olicable				_		
Test for overall effect: $Z = 2.3$:	2 (P = 0.02)						
03 Clomipramine vs Buspirone							
Pato 1991	9	21.22(5.49)	9	19.22(5.97)		-	→ 0.33 [-0.60, 1.26]
Subtotal (95% CI)	9		9				0.33 [-0.60, 1.26]
Test for heterogeneity: not app	olicable						
est for overall effect: Z = 0.7	0 (P = 0.48)						
otal (95% CI)	107		111				-0.22 [-0.49, 0.05]
est for heterogeneity: Chi² = 0 est for overall effect: Z = 1.60		= 0.18), I ² = 42.3%					•
					-1 -0.5	0 0.5	1
					Favours	SSRI Favours other dr	LIO.

Review: OCD adults: SSRIs

Comparison: 05 SSRIs/Clomipramine vs Non-SRIs (acute phase)

Outcome: 03 Adverse events



Review: OCD adults: SSRIs

Comparison: 05 SSRIs/Clomipramine vs Non-SRIs (acute phase)
Outcome: 04 Leaving study early due to adverse events

Study or sub-category	SSRI n/N	Other drug n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
D1 Sertraline vs Desipramine				
Hoehn-Saric 2000	8/80	22/86	-	0.39 [0.18, 0.83]
Subtotal (95% CI)	80	86	•	0.39 [0.18, 0.83]
Total events: 8 (SSRI), 22 (Other d	lrug)			
Test for heterogeneity: not applical				
est for overall effect: Z = 2.46 (P	= 0.01)			
2 Clomipramine vs Imipramine				
Volavka 1985	2/11	2/12		1.09 [0.18, 6.48]
Subtotal (95% CI)	11	12		1.09 [0.18, 6.48]
Total events: 2 (SSRI), 2 (Other dri				
Test for heterogeneity: not applical				
Γest for overall effect: Z = 0.10 (P	= 0.92)			
03 Fluvoxamine vs Desipramine				
Goodman 1990a	0/21	2/19		0.18 [0.01, 3.56]
Subtotal (95% CI)	21	19		0.18 [0.01, 3.56]
Total events: 0 (SSRI), 2 (Other dru				
Test for heterogeneity: not applica				
Test for overall effect: Z = 1.12 (P	= 0.26)			
04 Clomipramine vs Buspirone				
Pato 1991	1/10	0/10		3.00 [0.14, 65.90]
Subtotal (95% CI)	10	10		3.00 [0.14, 65.90]
otal events: 1 (SSRI), 0 (Other dr				
est for heterogeneity: not applical				
Test for overall effect: Z = 0.70 (P	= 0.49)			
05 Clomipramine vs Phenelzine				
Vallejo 1992	1/16	0/14		2.65 [0.12, 60.21]
Subtotal (95% CI)	16	14		2.65 [0.12, 60.21]
Total events: 1 (SSRI), 0 (Other dru				
[est for heterogeneity: not applical				
est for overall effect: Z = 0.61 (P	= 0.54)			
otal (95% CI)	138	141	•	0.51 [0.28, 0.95]
otal events: 12 (SSRI), 26 (Other				
est for heterogeneity: Chi² = 3.98		%		
est for overall effect: Z = 2.14 (P	= 0.03)			
		0.001	0.01 0.1 1 10 100	1000
			Favours SSRI Favours other o	drug

Review: OCD adults: SSRIs Comparison: 05 SSRIs/Clomipramine vs Non-SRIs (acute phase) Outcome: 05 Non-responders (OCD) SSRI Other drug RR (fixed) RR (fixed) Study or sub-category n/Ν n/N 95% CI 95% CI 02 Sertraline vs Desipramine Hoehn-Saric 2000 42/80 40/86 1.13 [0.83, 1.54] Subtotal (95% CI) 80 86 1.13 [0.83, 1.54] Total events: 42 (SSRI), 40 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.77 (P = 0.44) Total (95% CI) 80 86 1.13 [0.83, 1.54] Total events: 42 (SSRI), 40 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.77 (P = 0.44) 0.5 0.7 1.5 Favours SSRI Favours other drug OCD adults: SSRIs Review: Comparison: 06 SSRIs vs placebo (continuation phase) Outcome: 01 Leaving study early SSRI RR (fixed) RR (fixed) Study Placebo or sub-category n/N 95% CI 95% CI π/N 01 Fluoxetine Beasley 1992 3/23 0.39 [0.08, 1.84] 2/6 Montgomery 1993 16/61 2/12 1.57 [0.41, 5.97] Romano 2001 16/36 23/35 0.68 [0.44, 1.05] Subtotal (95% CD 0.75 [0.50, 1.12] 120 53 Total events: 35 (SSRI), 27 (Placebo) Test for heterogeneity: $Chi^2 = 2.07$, df = 2 (P = 0.36), $I^2 = 3.3\%$ Test for overall effect: Z = 1.40 (P = 0.16) 02 Paroxetine Hollander 2003d 19/53 0.50 [0.34, 0.75] 37/52 Subtotal (95% CI) 53 52 0.50 [0.34, 0.75] Total events: 19 (SSRI), 37 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 3.36 (P = 0.0008) Total (95% CI) 105 0.61 [0.46, 0.81] Total events: 54 (SSRI), 64 (Placebo) Test for heterogeneity: $Chi^2 = 3.36$, df = 3 (P = 0.34), $I^2 = 10.7\%$ Test for overall effect: Z = 3.39 (P = 0.0007) 0.01 0.1 100 10 Favours SSRI Favours placebo OCD adults: SSRIs Review: Comparison: 06 SSRIs vs placebo (continuation phase) Outcome: 02 Leaving study early due to adverse events RR (fixed) Study SSRI Placebo RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 Fluoxetine Montgomery 1993 2/61 0/12 1.05 [0.05, 20.58] Romano 2001 1.94 [0.18, 20.49] 2/36 1/35 Subtotal (95% CI) 47 1.54 [0.25, 9.58] Total events: 4 (SSRI), 1 (Placebo) Test for heterogeneity: $Chi^2 = 0.10^{\circ}$, df = 1 (P = 0.75), $I^2 = 0\%$ Test for overall effect: Z = 0.46 (P = 0.64) 02 Paroxetine Hollander 2003d 3/53 20/52 0.15 [0.05, 0.47] Subtotal (95% CI) 0.15 [0.05, 0.47] 53 52 Total events: 3 (SSRI), 20 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 3.26 (P = 0.001) Total (95% CI) 150 99 0.26 [0.11, 0.63] Total events: 7 (SSRI), 21 (Placebo) Test for heterogeneity: Chi² = 4.58, df = 2 (P = 0.10), I² = 56.3% Test for overall effect: Z = 3.00 (P = 0.003)0.01 0.1 10 100

Favours SSRI Favours placebo

Review: OCD adults: SSRIs Comparison: 06 SSRIs vs placebo (continuation phase) Outcome: 03 Non-responders SSRI Placebo RR (fixed) RR (fixed) Study 95% CI or sub-category nNn/N 95% CI 01 Fluoxetine Beasley 1992 14/23 3/6 1.22 [0.51, 2.89] Subtotal (95% CI) 23 6 1.22 [0.51, 2.89] Total events: 14 (SSRI), 3 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.45 (P = 0.66) 0.2 0.5 2 5 Favours SSRI Favours placebo Review OCD adults: SSRIs 06 SSRIs vs placebo (continuation phase) Comparison: 04 Y-BOCS Outcome: Study SSRI Placebo SMD (fixed) SMD (fixed) or sub-category Mean (SD) Ν Mean (SD) Ν 95% CI 95% CI 01 Fluoxetine Beasley 1992 23 9.80(7.30) 6 8.50(6.70) 0.18 [-0.72, 1.08] Romano 2001 35 1.46(7.08) 34 2.50(6.16) -0.15 [-0.63, 0.32] -0.08 [-0.50, 0.34] Subtotal (95% CI) 58 40 Test for heterogeneity: Chi² = 0.41, df = 1 (P = 0.52), l² = 0% Test for overall effect: Z = 0.39 (P = 0.70) -4 0 Favours SSRI Favours placebo Review: OCD adults: SSRIs 06 SSRIs vs placebo (continuation phase) Comparison: Outcome: 05 Hamilton Rating Scale for Depression Study SSRI Placebo SMD (fixed) SMD (fixed) or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Romano 2001 35 1.20(5.37) 34 1.79(5.06) -0.11 [-0.58, 0.36] Subtotal (95% CI) 35 34 -0.11 [-0.58, 0.36] Test for heterogeneity; not applicable Test for overall effect: Z = 0.46 (P = 0.64) -1 -0.5Ó 0.5 Favours SSRI Favours placebo OCD adults: SSRIs Review: Comparison: 06 SSRIs vs placebo (continuation phase) Outcome 06 SF-36; social functioning SMD (fixed) SMD (fixed) Study SSRI Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Romano 2001 0.15 [-0.32, 0.63] 34 9.56(19.23) 34 5.88(27.22) Subtotal (95% CI) 34 0.15 [-0.32, 0.63] Test for heterogeneity: not applicable Test for overall effect: Z = 0.64 (P = 0.53) -0.5 Ó 0.5 Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 07 SSRIs vs placebo (discontinuation phase) Outcome: 01 Death SSRI RR (fixed) RR (fixed) Study Placebo or sub-category n/Ν n/N 95% CI 95% CI 01 Sertraline Koran 2002 0/109 1/114 0.35 [0.01, 8.46] Subtotal (95% CI) 0.35 [0.01, 8.46] 109 114 Total events: 0 (SSRI), 1 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.65 (P = 0.52)

0.01

0.1

Favours SSRI

10

Favours placebo

100

Review: OCD adults: SSRIs 07 SSRIs vs placebo (discontinuation phase) Comparison: Outcome: 02 Leaving study early Study SSRI Placebo RR (fixed) RR (fixed) n/N n/N 95% CI 95% CI or sub-category 01 Sertraline 0.58 [0.42, 0.82] 0.58 [0.42, 0.82] Koran 2002 33/109 59/114 Subtotal (95% CI) 109 114 Total events: 33 (SSRI), 59 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 3.13 (P = 0.002) 0.2 0.5 ż Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 07 SSRIs vs placebo (discontinuation phase) Outcome: 03 Leaving study early due to adverse effects Study SSRI RR (fixed) RR (fixed) Placebo or sub-category n/N n/N 95% CI 95% CI 01 Sertraline Koran 2002 5/109 12/114 0.44 [0.16, 1.20] Subtotal (95% CI) 0.44 [0.16, 1.20] 109 114 Total events: 5 (SSRI), 12 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.61 (P = 0.11) 0.2 0.5 5 10 Favours SSRI Favours placebo OCD adults: SSRIs Review: 07 SSRIs vs placebo (discontinuation phase) Comparison: Outcome: 04 Relapse SSRI Placebo RR (fixed) RR (fixed) Study n/N n/N 95% CI 95% CI or sub-category 01 Sertraline Koran 2002 4/109 6/114 0.70 [0.20, 2.40] Subtotal (95% CI) 0.70 [0.20, 2.40] Total events: 4 (SSRI), 6 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.57 (P = 0.57) 0.2 Ś 0.5 2 Favours SSRI Favours placebo Review: OCD adults: SSRIs Comparison: 07 SSRIs vs placebo (discontinuation phase) 05 Y-BOCS

Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Placebo Mean (SD)	SMD (959	,	SMD (fixed) 95% Cl
01 Sertraline							
Koran 2002	108	11.40(7.50)	113	14.20(7.70)			-0.37 [-0.63, -0.10]
Subtotal (95% CI)	108		113				-0.37 [-0.63, -0.10]
Test for heterogeneity: not	applicable				_		
Test for overall effect: Z =	2.70 (P = 0.007)						
					-1 -0.5 0	0.5	;

Favours SSRI Favours placebo

OCD adults: SSRIs Review:

Comparison: 07 SSRIs vs placebo (discontinuation phase)

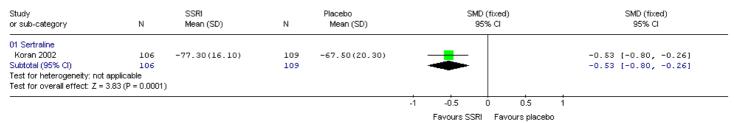
06 NIMHLOC Outcome

Study or sub-category	N	SSRI Mean (SD)	N	Placebo Mean (SD)		SMD (fixe 95% C		SMD (fixed) 95% Cl
01 Sertraline Koran 2002 Subtotal (95% CI) Test for heterogeneity: not Test for overall effect: Z =		4.80(2.60)	113 113	5.70(2.70)		+		-0.34 [-0.60, -0.07] -0.34 [-0.60, -0.07]
					-1	-0.5 0	0.5	1
						Favours SSRI F	avours placebo	

Review: OCD adults: SSRIs

Comparison: 07 SSRIs vs placebo (discontinuation phase)

07 QoL Enjoyment and Satisfaction Outcome:



Review: OCD adults: SSRIs

08 SSRIs at different doses (continuation phase) Comparison:

Outcome: 01 Leaving study early

Study ir sub-category	Dose 1 n/N	Dose 2 n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
11 40mg Fluoxetine v 20mg Fluo:	xetine			
Beasley 1992	6/21	3/23		- 2.19 [0.63, 7.67]
Montgomery 1993	6/20	5/18		1.08 [0.40, 2.94]
Subtotal (95% CI)	41	41	-	1.47 [0.68, 3.19]
otal events: 12 (Dose 1), 8 (Dos	se 2)			
est for heterogeneity: Chi² = 0.7	75, df = 1 (P = 0.39), l² = 09	6		
est for overall effect: Z = 0.98 ((P = 0.33)			
2 60mg Fluoxetine v 20mg Fluo:	xetine			
Beasley 1992	3/26	3/23		0.88 [0.20, 3.96]
Montgomery 1993	5/23	5/18		0.78 [0.27, 2.29]
Subtotal (95% CI)	49	41		0.82 [0.34, 1.97]
otal events: 8 (Dose 1), 8 (Dose	e 2)		-	
est for heterogeneity: Chi² = 0.0	02 , df = 1 (P = 0.90), I^2 = 0.90	6		
est for overall effect: Z = 0.45 ((P = 0.66)			
3 60mg Fluoxetine v 40mg Fluo:	xetine			
Beasley 1992	3/26	6/21 -		0.40 [0.11, 1.43]
Montgomery 1993	5/23	6/20		0.72 [0.26, 2.02]
Subtotal (95% CI)	49	41	-	0.56 [0.26, 1.24]
otal events: 8 (Dose 1), 12 (Dos	se 2)		-	
est for heterogeneity: Chi² = 0.5	50 , df = 1 (P = 0.48), I^2 = 0.9	6		
est for overall effect: Z = 1.43 ((P = 0.15)			

Favours Dose 1 Favours Dose 2

Review: OCD adults: SSRIs

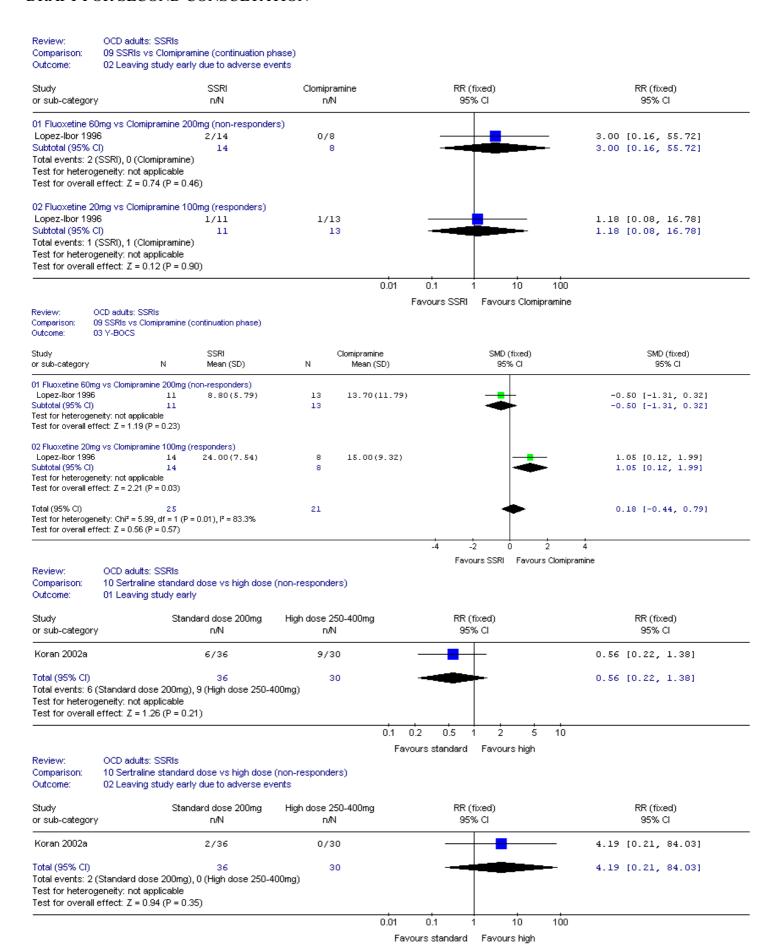
Comparison: 08 SSRIs at different doses (continuation phase) Outcome: 02 Leaving study early due to adverse events

Study or sub-category	Dose 1 n/N	Dose 2 n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl		
01 40mg Fluoxetine v 20mg Fluoxe	etine					
Montgomery 1993	1/20	0/18		2.71 [0.12, 62.70]		
Subtotal (95% CI)	20	18		2.71 [0.12, 62.70]		
Total events: 1 (Dose 1), 0 (Dose	2)					
Test for heterogeneity: not applica	able					
Test for overall effect: Z = 0.62 (P	P = 0.53)					
02 60mg Fluoxetine v 20mg Fluoxe	etine					
Montgomery 1993	1/23	0/18		2.38 [0.10, 55.06]		
Subtotal (95% CI)	23	18		2.38 [0.10, 55.06]		
Total events: 1 (Dose 1), 0 (Dose	2)					
Test for heterogeneity: not applica	able					
Test for overall effect: Z = 0.54 (P	P = 0.59)					
03 60mg Fluoxetine v 40mg Fluoxe	etine					
Montgomery 1993	1/23	1/20		0.87 [0.06, 13.02]		
Subtotal (95% CI)	23	20		0.87 [0.06, 13.02]		
Total events: 1 (Dose 1), 1 (Dose	2)					
Test for heterogeneity: not applica	able					
Test for overall effect: Z = 0.10 (P	P = 0.92)					
		0.	01 0.1 1 10	100		
		Favours Dose 1 Favours Dose 2				

Review: OCD adults: SSRIs Comparison: 08 SSRIs at different doses (continuation phase) Outcome: 03 Non-responders Dose 1 Dose 2 RR (fixed) RR (fixed) Study or sub-category n/N n/N 95% CI 95% CI 01 Fluoxetine 40mg v Fluoxetine 20mg Beasley 1992 12/21 14/23 0.94 [0.57, 1.54] Subtotal (95% CI) 23 0.94 [0.57, 1.54] Total events: 12 (Dose 1), 14 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 0.25 (P = 0.80) 02 Fluoxetine 60mg v Fluoxetine 20mg Beasley 1992 11/26 14/23 0.70 [0.40, 1.21] Subtotal (95% CI) 0.70 [0.40, 1.21] 23 Total events: 11 (Dose 1), 14 (Dose 2) Test for heterogeneity: not applicable Test for overall effect: Z = 1.28 (P = 0.20) 03 Fluoxetine 60mg v Fluoxetine 40mg Beasley 1992 11/26 12/21 0.74 [0.41, 1.32] Subtotal (95% CI) 0.74 [0.41, 1.32] 26 Total events: 11 (Dose 1), 12 (Dose 2) Test for heterogeneity; not applicable Test for overall effect: Z = 1.01 (P = 0.31) 0.2 0.5 ż 5 Favours Dose 1 Favours Dose 2 Review: OCD adults: SSRIs Comparison: 08 SSRIs at different doses (continuation phase) 04 Y-BOCS Outcome: SMD (fixed) SMD (fixed) Dose 2 or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CL 01 Fluoxetine 40mg v Fluoxetine 20mg Beasley 1992 9.40(6.00) 23 9.80(7.30) -0.06 [-0.65, 0.53] Subtotal (95% CI) 21 23 -0.06 [-0.65, 0.53] Test for heterogeneity; not applicable Test for overall effect: Z = 0.19 (P = 0.85) 02 Fluoxetine 60mg v Fluoxetine 20mg Beasley 1992 26 7.00(5.40) 23 9.80(7.30) -0.43 [-1.00, 0.13] Subtotal (95% CI) -0.43 [-1.00, 0.13] 26 23 Test for heterogeneity: not applicable Test for overall effect: Z = 1.49 (P = 0.14) 03 Fluoxetine 60mg v Fluoxetine 40mg Beasley 1992 26 7.00(5.40) 21 9.40(6.00) -0.42 [-1.00, 0.17] Subtotal (95% CI) -0.42 [-1.00, 0.17] Test for heterogeneity: not applicable Test for overall effect: Z = 1.40 (P = 0.16) Ó Favours Dose 1 Favours Dose 2 OCD adults: SSRIs Review: Comparison: 09 SSRIs v Clomipramine (continuation phase) Outcome: 01 Leaving study early Study SSRI Clomipramine RR (fixed) RR (fixed) or sub-category n/N n/N 95% CI 95% CI 01 Fluoxetine 60mg vs Clomipramine 200mg (non-responders) Lopez-Ibor 1996 3/8 1.14 [0.39, 3.36] 6/14 Subtotal (95% CI) 1.14 [0.39, 3.36] Total events: 6 (SSRI), 3 (Clomipramine) Test for heterogeneity: not applicable Test for overall effect: Z = 0.24 (P = 0.81) 02 Fluoxetine 20mg vs Clomipramine 100mg (responders) Lopez-Ibor 1996 0.59 [0.19, 1.83] 3/11 6/13 Subtotal (95% CI) 0.59 [0.19, 1.83] 11 13 Total events: 3 (SSRI), 6 (Clomipramine) Test for heterogeneity: not applicable Test for overall effect: Z = 0.91 (P = 0.36) 0.1 0.2 0.5 ż 5 10

Favours SSRI

Favours Clomipramine



Review: OCD adults: SSRIs Comparison: 10 Sertraline standard dose vs high dose (non-responders) Outcome: 03 Non-responders Standard dose 200mg High dose 250-400mg RR (fixed) RR (fixed) 95% CI or sub-category n/N n/N 95% CI Koran 2002a 24/36 18/30 1.11 [0.77, 1.61] Total (95% CI) 36 30 1.11 [0.77, 1.61] Total events: 24 (Standard dose 200mg), 18 (High dose 250-400mg) Test for heterogeneity: not applicable Test for overall effect: Z = 0.55 (P = 0.58) 0.1 0.2 0.5 5 10 Favours standard Favours high Review: OCD adults: SSRIs 11 SSRI vs Placebo (maintenance) Comparison: 01 Leaving the study early Outcome: Study SSRI RR (fixed) RR (fixed) Placebo or sub-category n/N n/N 95% CI 95% CI 01 Paroxetine Ansseau 1995 0.64 [0.35, 1.16] 19/51 7/12 Subtotal (95% CI) 51 12 0.64 [0.35, 1.16] Total events: 19 (SSRI), 7 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.47 (P = 0.14) 0.2 0.5 5 10 Favours SSRI Favours Placebo OCD adults: SSRIs Review: Comparison: 11 SSRI vs Placebo (maintenance) Outcome: 02 Leaving the study early due to adverse events SSRI RR (fixed) RR (fixed) Study Placebo n/N 95% CI 95% CI or sub-category n/N 01 Paroxetine Ansseau 1995 8/51 2/12 0.94 [0.23, 3.88] Subtotal (95% CI) 0.94 [0.23, 3.88] 51 Total events: 8 (SSRI), 2 (Placebo) Test for heterogeneity; not applicable Test for overall effect: Z = 0.08 (P = 0.93) 10 5 0.1 0.2 0.5 2 Favours SSRI Favours Placebo OCD adults: SSRIs Review: Comparison: 11 SSRI vs Placebo (maintenance) Outcome: 03 Adverse events Study SSRI Placebo RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 Paroxetine Ansseau 1995 1.24 [0.81, 1.88] 42/51 8/12 Subtotal (95% CI) 51 12 1.24 [0.81, 1.88] Total events: 42 (SSRI), 8 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.99 (P = 0.32) 0.2 0.5 Favours SSRI Favours Placebo OCD adults: SSRIs Review: Comparison: 11 SSRI vs Placebo (maintenance) Outcome: 04 Severe adverse events SSRI RR (fixed) RR (fixed) Study Placebo or sub-category n/N 95% CI 95% CI n/N 01 Paroxetine Ansseau 1995 18/51 3/12 1.41 [0.49, 4.03] Subtotal (95% CI) 12 1.41 [0.49, 4.03] 51 Total events: 18 (SSRI), 3 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.64 (P = 0.52)

0.1 0.2

0.5

Favours SSRI

ż

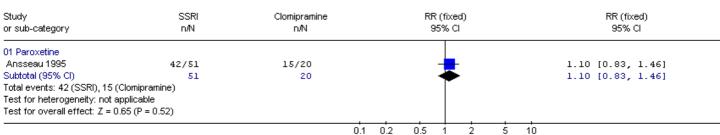
Favours Placebo

5 10

Review: OCD adults: SSRIs Comparison: 11 SSRI vs Placebo (maintenance) Outcome: 05 Non-responders (Y-BOCS 25% reduction) Study SSRI Placebo RR (fixed) RR (fixed) 95% CI 95% CI or sub-category n/N 01 Paroxetine Ansseau 1995 0.63 [0.20, 2.02] 8/51 3/12 Subtotal (95% CI) 51 12 0.63 [0.20, 2.02] Total events: 8 (SSRI), 3 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.78 (P = 0.43) 0.1 0.2 0.5 5 10 Favours SSRI Favours Placebo Review OCD adults: SSRIs 11 SSRI vs Placebo (maintenance) Comparison: 06 Y-BOCS Outcome: Study SSRI Placebo SMD (fixed) SMD (fixed) N Mean (SD) Ν Mean (SD) or sub-category 95% CI 95% CI 01 Paroxetine Ansseau 1995 51 -14.70(7.20) 12 -11.00(6.90) -0.51 [-1.15, 0.12] Subtotal (95% CI) 51 12 -0.51 [-1.15, 0.12] Test for heterogeneity; not applicable Test for overall effect: Z = 1.58 (P = 0.11) -10 -5 ń 10 Favours SSRI Favours Placebo OCD adults: SSRIs Comparison: 11 SSRI vs Placebo (maintenance) 07 NIMH-OCS Outcome: Study SSRI Placebo SMD (fixed) SMD (fixed) or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Paroxetine Ansseau 1995 -0.61 [-1.25, 0.03] 51 -5.20(2.60) 12 -3.60(2.60) Subtotal (95% CI) 12 -0.61 [-1.25, 0.03] Test for heterogeneity: not applicable Test for overall effect: Z = 1.87 (P = 0.06) -10 -5 Ó 10 Favours SSRI Favours Placebo Review: OCD adults: SSRIs Comparison: 12 SSRI vs Clomipramine (maintenance) Outcome: 01 Leaving the study early SSRI RR (fixed) RR (fixed) Study Clomipramine or sub-category n/N 95% CI 95% CI n/N 01 Paroxetine Ansseau 1995 19/51 8/20 0.93 [0.49, 1.77] Subtotal (95% CI) 20 0.93 [0.49, 1.77] Total events: 19 (SSRI), 8 (Clomipramine) Test for heterogeneity: not applicable Test for overall effect: Z = 0.22 (P = 0.83) 10 0.1 0.2 0.5 2 5 Favours SSRI Favours Clomipramine Review: OCD adults: SSRIs 12 SSRI vs Clomipramine (maintenance) Comparison: Outcome: 02 Leaving the study early due to adverse events

Study or sub-category	SSRI n/N	Clomipramine n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
O1 Paroxetine Ansseau 1995 Subtotal (95% CI) Total events: 8 (SSRI), 6 (Clor Test for heterogeneity: not ap Test for overall effect: Z = 1.3	plicable	6/20 20		0.52 [0.21, 1.32] 0.52 [0.21, 1.32]
		0	.1 0.2 0.5 1 2 5 Favours SSRI Favours Clomipra	10 mine

Review: OCD adults: SSRIs 12 SSRI vs Clomipramine (maintenance) Comparison: Outcome: 03 Adverse events



Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 12 SSRI vs Clomipramine (maintenance)

Outcome: 04 Severe adverse events

Study SSRI RR (fixed) RR (fixed) Clomipramine 95% CI or sub-category n/N n/N 95% CI 01 Paroxetine Ansseau 1995 18/51 2/20 3.53 [0.90, 13.84] Subtotal (95% CI) 51 20 3.53 [0.90, 13.84] Total events: 18 (SSRI), 2 (Clomipramine) Test for heterogeneity: not applicable Test for overall effect: Z = 1.81 (P = 0.07) 0.01 0.1 100

Favours SSRI Favours Clomipramine

OCD adults: SSRIs Review:

12 SSRI vs Clomipramine (maintenance) Comparison: Outcome: 05 Responders (Y-BOCS 25% reduction)

3/20		1.05 [0.31, 3.55]
20		1.05 [0.31, 3.55]
	T	·
_	4	01 02 05 1 2 5 1

Favours SSRI Favours Clomipramine

Favours SSRI Favours Clomipramine

Review: OCD adults: SSRIs

Comparison: 12 SSRI vs Clomipramine (maintenance)

06 Y-BOCS Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)		MD (fixed) 95% CI		SMD (fixed) 95% Cl
01 Paroxetine								
Ansseau 1995	51	-14.70(7.20)	20	-14.60(7.10)				-0.01 [-0.53, 0.50]
Subtotal (95% CI)	51		20			•		-0.01 [-0.53, 0.50]
Test for heterogeneity: not ap	plicable					I		
Test for overall effect: $Z = 0.0$	05 (P = 0.96)							

OCD adults: SSRIs Review:

Comparison: 12 SSRI vs Clomipramine (maintenance)

07 NIMH-OCS Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Clomipramine Mean (SD)		:	SMD (fixed 95% CI	1)		SMD (fixed) 95% Cl
01 Paroxetine										
Ansseau 1995	51	-5.20(2.60)	19	-4.20(2.20)						-0.40 [-0.93, 0.14]
Subtotal (95% CI)	51		19				- 4			-0.40 [-0.93, 0.14]
Test for heterogeneity: not	t applicable						1			
Test for overall effect: Z =	1.46 (P = 0.14)									
					-10	-5	-	5	10	
						Favours S	SSRI Fa	vours Clor	nipramine	

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 01 Leaving the study early

Study or sub-category	Paroxetine n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
Ansseau 1995	0/14	3/18 -		0.18 [0.01, 3.24]
Subtotal (95% CI)	14	18 -		0.18 [0.01, 3.24]
Total events: 0 (Paroxetine), 3 i				
Test for heterogeneity: not app Test for overall effect: Z = 1.16				
02 Paroxetine at 6 months				
Bailer 1995	9/20	16/24	-	0.68 [0.39, 1.18]
Subtotal (95% CI)	20	24	•	0.68 [0.39, 1.18]
Total events: 9 (Paroxetine), 16	(Placebo)		-	
Test for heterogeneity: not app	licable			
Test for overall effect: Z = 1.37	(P = 0.17)			
Total (95% CI)	34	42	•	0.59 [0.33, 1.04]
Total events: 9 (Paroxetine), 19	(Placebo)			
Test for heterogeneity: Chi² = 0	1.87 , df = 1 (P = 0.35), $I^2 = 0\%$	6		
Test for overall effect: $Z = 1.83$	3 (P = 0.07)			
		0.0	1 0.1 1 10	100
		F	avours Paroxetine Favours Place	bo

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)
Outcome: 02 Leaving the study early due to adverse events

Paroxetine Placebo RR (fixed) RR (fixed) Study 95% CI or sub-category nΝ n/N 95% CI 01 Paroxetine at 6 months Bailer 1995 0/20 9/24 0.06 [0.00, 1.01] Subtotal (95% CI) 20 24 0.06 [0.00, 1.01] Total events: 0 (Paroxetine), 9 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.95 (P = 0.05) 0.001 0.01 0.1 10 100 1000

Favours Paroxetine

Favours Placebo

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

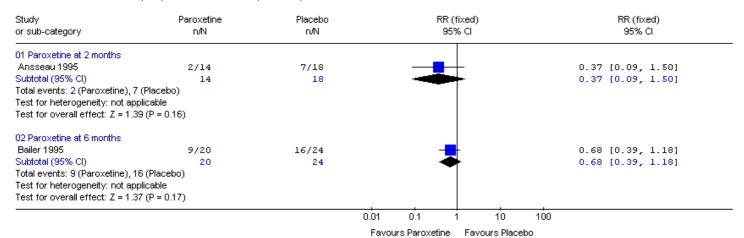
Outcome: 03 Full relapse (Y-BOCS and CGI Severity increase)

Study or sub-category	Paroxetine n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% CI
01 Paroxetine at 6 months				
Bailer 1995	1/20	6/24		0.20 [0.03, 1.53]
Subtotal (95% CI)	20	24		0.20 [0.03, 1.53]
Total events: 1 (Paroxetine),	6 (Placebo)		 -	
Test for heterogeneity: not a	pplicable			
Test for overall effect: Z = 1	.55 (P = 0.12)			
		0.0	1 0.1 1 10	100
		0.0	1 0.1 1 10	100
		F	avours Paroxetine Favours Place	bo

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 04 Partial relapse (Y-BOCS or CGI Severity increase)



Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 05 Adverse events

Study or sub-category	Paroxetine n/N	Placebo n/N		RR (fixed) 95% Cl		RR (fixed) 95% Cl
01 Paroxetine at 2 months						
Ansseau 1995	8/14	9/18				1.14 [0.60, 2.18]
Subtotal (95% CI)	14	18				1.14 [0.60, 2.18]
Total events: 8 (Paroxetine),	9 (Placebo)			_		
Test for heterogeneity; not a	applicable					
Test for overall effect: Z = 0	.40 (P = 0.69)					
	· · · · · · · · · · · · · · · · · · ·					
			0.1 0.2	0.5 1 2	5 10	

Favours Paroxetine

Favours Placebo

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 06 Severe adverse events

Study or sub-category	Paroxetine n/N	Placebo n <i>i</i> N	RR (fixed) 95% Cl		RR (fixed) 95% Cl
01 Paroxetine at 2 months					
Ansseau 1995	1/14	4/18			0.32 [0.04, 2.57]
Subtotal (95% CI)	14	18			0.32 [0.04, 2.57]
Total events: 1 (Paroxetine),	4 (Placebo)		- <u>-</u> -		
Test for heterogeneity: not a	pplicable				
Test for overall effect: Z = 1	.07 (P = 0.28)				
			0.01 0.1 1	10 100	
			Favours Paroxetine Fav	ours Placebo	

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 07 Y-BOCS

Study or sub-category	N	Paroxetine Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Paroxetine at 2 months						
Ansseau 1995	12	0.80(1.60)	12	0.90(3.20)	-	-0.04 [-0.84, 0.76]
Subtotal (95% CI)	12		12		•	-0.04 [-0.84, 0.76]
Test for heterogeneity: not a	applicable				Ī	
Test for overall effect: $Z = 0$	0.09 (P = 0.93)					
02 Paroxetine at 6 months						
Bailer 1995	12	-3.50(3.60)	8	0.88(3.59)	-	-1.17 [-2.15, -0.19]
Subtotal (95% CI)	12		8		•	-1.17 [-2.15, -0.19]
Test for heterogeneity: not a	applicable				T	
Test for overall effect: $Z = 2$	2.33 (P = 0.02)					
Total (95% CI)	24		20		•	-0.49 [-1.11, 0.13]
Test for heterogeneity: Chi ²	= 3.05, df = 1 (P	$= 0.08$), $I^2 = 67.2\%$			1	
Test for overall effect: Z = 1		••				
				-1	0 -5 0 5	10

Favours Paroxetine Favours Placebo

Review: OCD adults: SSRIs

Comparison: 13 SSRI vs Placebo (relapse prevention)

Outcome: 08 NIMH-OC

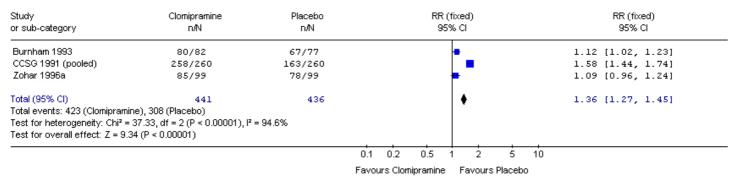
Study or sub-category	N	Paroxetine Mean (SD)	N	Placebo Mean (SD)		SMD (fixed) 95% Cl		SMD (fixed) 95% Cl
01 Paroxetine at 2 months								
Ansseau 1995	11	0.20(0.90)	9	1.30(2.30)		-		-0.63 [-1.54, 0.28]
Subtotal (95% CI)	11		9			•		-0.63 [-1.54, 0.28]
Test for heterogeneity: not ap	plicable							
Test for overall effect: $Z = 1.3$	36 (P = 0.17)							
02 Paroxetine at 6 months								
Bailer 1995	12	-0.67(1.66)	8	0.38(1.67)		-		-0.60 [-1.52, 0.31]
Subtotal (95% CI)	12		8			•		-0.60 [-1.52, 0.31]
Test for heterogeneity: not ap	plicable							
Test for overall effect: $Z = 1.2$	29 (P = 0.20)							
Total (95% CI)	23		17			•		-0.62 [-1.26, 0.03]
Test for heterogeneity: Chi ² =	0.00, df = 1 (P	= 0.97), I ² = 0%						
Test for overall effect: $Z = 1.8$	B7 (P = 0.06)							
				-	-10	5 0	5 10	
					Favours Pa	roxetine Favou	urs Placebo	

Clomipramine (OCD, adults)

Review: OCD adults: Clomipramine

Comparison: 01 Clomipramine vs placebo: acute phase

Outcome: 01 Adverse events



Review: OCD adults: Clomipramine

01 Clomipramine vs placebo: acute phase Comparison:

Outcome: 02 Leaving study early

Study or sub-category	Clomipramine n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% CI
Burnham 1993	28/82	20/77	-	1.31 [0.81, 2.13]
CCSG 1991(1)	17/118	13/121	- •	1.34 [0.68, 2.64]
CCSG 1991(2)	14/142	12/139	 	1.14 [0.55, 2.38]
Stein 1992	7/21	2/23		→ 3.83 [0.89, 16.44]
Zohar 1996a	34/99	39/99	-	0.87 [0.60, 1.26]
Total (95% CI)	462	459	◆	1.15 [0.90, 1.47]
Total events: 100 (Clomipra	7			
Test for heterogeneity: Chi ²	' = 5.32, df = 4 (P = 0.26), l² = 24.5	9%		
Test for overall effect: Z = :	1.11 (P = 0.27)			
		0.	1 0.2 0.5 1 2 5	10

OCD adults: Clomipramine Review:

Comparison: 01 Clomipramine vs placebo; acute phase Outcome: 03 Leaving study early due to adverse events

Study or sub-category	Clomipramine n/N	Placebo n/N	RR (f 95%	,	RR (fixed) 95% Cl
Burnham 1993	17/82	8/77	-		2.00 [0.91, 4.36]
Zohar 1996a	17/99	6/99			2.83 [1.17, 6.89]
Total (95% CI)	181	176		-	2.35 [1.31, 4.22]
Total events: 34 (Clomipram	nine), 14 (Placebo)			_	
Test for heterogeneity: Chi ²	$= 0.34$, df = 1 (P = 0.56), $I^2 = 0\%$				
Test for overall effect: Z = 2	2.86 (P = 0.004)				
			0.1 0.2 0.5 1	2 5 10	

Favours Clomipramine Favours Placebo

OCD adults: Clomipramine Review:

01 Clomipramine vs placebo: acute phase Comparison:

Outcome: 04 Non-responders (Y-BOCS)

Study or sub-category	Clomipramine n/N	Placebo n/N				R (fixed 95% CI				RR (fixed) 95% Cl
Burnham 1993	42/82	45/77			_	_				0.88 [0.66, 1.16]
Zohar 1996a	47/99	64/99			-	-				0.73 [0.57, 0.95]
Total (95% CI)	181	176			4	•				0.79 [0.66, 0.96]
Total events: 89 (Clomipram	ine), 109 (Placebo)									
Test for heterogeneity: Chi²	$= 0.83$, df = 1 (P = 0.36), $I^2 = 0\%$									
Test for overall effect: Z = 2	2.40 (P = 0.02)									
			0.1	0.2	0.5	1	ż	5	10	

Favours Clomipramine Favours Placebo

Favours Clomipramine Favours Placebo

OCD adults: Clomipramine Comparison:

01 Clomipramine vs placebo: acute phase

Outcome: 05 Non-responders (CGI)

Study or sub-category	Clomipramine n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
Stein 1992	14/21	17/23	-	0.90 [0.61, 1.33]
Total (95% CI) Total events: 14 (Clomipran Test for heterogeneity: not Test for overall effect: Z =	applicable	23	+	0.90 [0.61, 1.33]
		0.1	0.2 0.5 1 2	5 10
		Favor	urs Clomipramine Favours F	Placeho

Review: OCD adults: Clomipramine

Comparison: 01 Clomipramine vs placebo: acute phase

Outcome: 06 Non-responders (all)

Study or sub-category	Clomipramine n/N	Placebo n/N			,	(fixed) % Cl			RR (fixed) 95% Cl
Burnham 1993	42/82	45/77			-	-			0.88 [0.66, 1.16]
Stein 1992	14/21	17/23			-	<u> </u>			0.90 [0.61, 1.33]
Zohar 1996a	47/99	64/99			-	-			0.73 [0.57, 0.95]
Total (95% CI)	202	199			•				0.81 [0.68, 0.96]
Total events: 103 (Clomipra	mine), 126 (Placebo)				•				•
Test for heterogeneity: Chi ²	= 1.17, df = 2 (P = 0.56), l ² = 0%								
Test for overall effect: Z = :									
			0.1	0.2	0.5	1 2	5	10	

Favours Clomipramine Favours placebo

Review: OCD adults: Clomipramine

Comparison: 01 Clomipramine vs placebo: acute phase

Outcome: 07 Non-remitters

Study or sub-category	Clomipramine n/N	Placebo n/N			fixed) % Cl		RR (fixed) 95% Cl
CCSG 1991 (pooled)	136/260	252/260					0.54 [0.48, 0.61]
Total (95% CI) Total events: 136 (Clomipramir Test for heterogeneity: not ap Test for overall effect: Z = 10	plicable	260		•			0.54 [0.48, 0.61]
			0.1 0.2	0.5	1 2	5 10	

Favours Clomipramine Favours placebo

Review: OCD adults: Clomipramine

Comparison: 01 Clomipramine vs placebo: acute phase

Outcome: 08 Y-BOCS

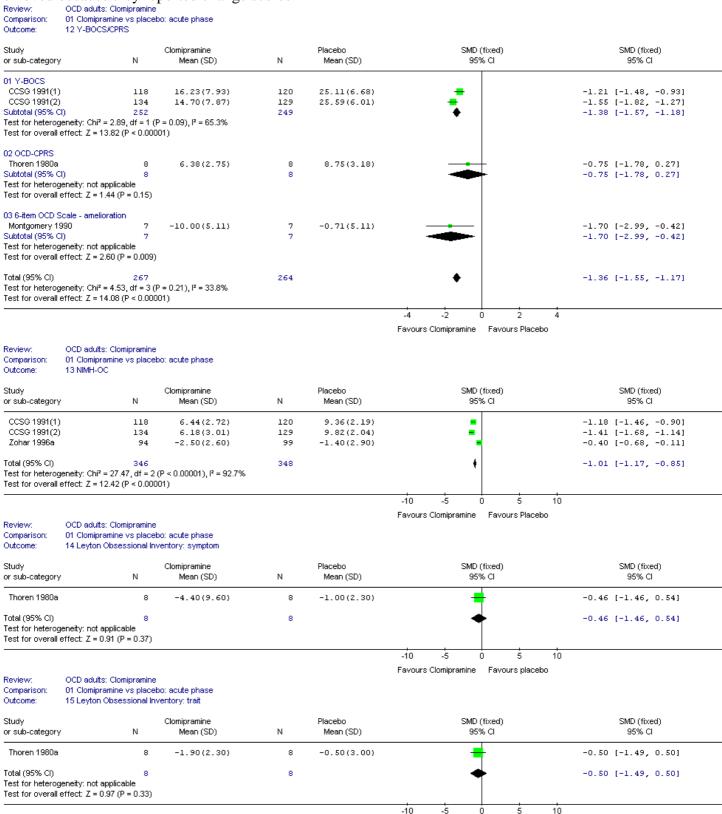
Study or sub-category	N	Clomipramine Mean (SD)	N	Placebo Mean (SD)			MD (fixed) 95% Cl)		SMD (fixed) 95% CI
Burnham 1993	78	-7.73(7.42)	75	-4.60(7.53)			-			-0.42 [-0.74, -0.10]
CCSG 1991(1)	118	16.23(7.93)	120	25.11(6.68)		-				-1.21 [-1.48, -0.93]
CCSG 1991(2)	134	14.70(7.87)	129	25.59(6.01)		-				-1.55 [-1.82, -1.27]
Zohar 1996a	94	-8.00(8.20)	99	-5.00(7.90)			-			-0.37 [-0.66, -0.09]
Total (95% CI)	424		423			•	,			-0.93 [-1.07, -0.78]
Test for heterogeneity: Ch	$i^2 = 47.71$, $df = 3$	(P < 0.00001), I ² = 93.7%				•				•
Test for overall effect: Z =	= 12.62 (P < 0.0000	01)								
					-4	-2	0	2	4	

Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo: acute phase Outcome 09 OCD Scale (CPRS) SMD (fixed) SMD (fixed) Study Clomipramine Placebo or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI Thoren 1980a 8 6.38(2.75) 8 8.75(3.18) -0.75 [-1.78. 0.27] Total (95% CI) -0.75 [-1.78, 0.27] Test for heterogeneity: not applicable Test for overall effect: Z = 1.44 (P = 0.15) -10 -5 ò 10 Favours Clomipramine Favours Placebo OCD adults: Clomipramine Review: Comparison: 01 Clomipramine vs placebo: acute phase Outcome 10 6-item OCD Scale - amelioration SMD (fixed) SMD (fixed) Study Placebo Clomipramine Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Montgomery 1990 -10.00(5.11) -0.71(5.11) -1.70 [-2.99, -0.42] Total (95% CI) -1.70 [-2.99] -0.421 Test for heterogeneity: not applicable Test for overall effect: Z = 2.60 (P = 0.009) -10 -5 10 0 Favours Clomipramine Favours placebo Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo; acute phase 11 OCD (OCD-CPRS; 6-item OCD Scale-amelioration) Outcome Study Clomipramine Placebo SMD (fixed) SMD (fixed) Ν Ν or sub-category Mean (SD) Mean (SD) 95% CI 95% CI Montgomery 1990 -10.00(5.11) -0.71(5.11) -1.70 [-2.99, -0.42] Thoren 1980a 8 6.38(2.75) 8 8.75(3.18) -0.75 [-1.78, 0.27] Total (95% CI) 15 -1.12 [-1.92, -0.32] Test for heterogeneity: Chi² = 1.28, df = 1 (P = 0.26), l² = 21.8% Test for overall effect: Z = 2.75 (P = 0.006) 10 -10 -5 0 Favours Clomipramine Favours placebo Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo: acute phase 12 Y-BOCS/CPRS Outcome Clomipramine Placebo SMD (fixed) SMD (fixed) or sub-category M Mean (SD) M Mean (SD) 95% CI 95% CI 01 Y-BOCS Burnham 1993 4.60(7.53) -0.42 [-0.74, -0.10] -1.21 [-1.48, -0.93] -1.55 [-1.82, -1.27] CCSG 1991(1) 118 16.23(7.93) 120 25.11(6.68) CCSG 1991(2) 134 14.70(7.87) 129 25.59(6.01) Zohar 1996a -0.37 [-0.66, -0.09] 94 -8.00(8.20) 99 -5.00(7.90) Subtotal (95% CI) -0.93 [-1.07, -0.78] Test for heterogeneity: Chi² = 47.71, df = 3 (P < 0.00001), l² = 93.7% Test for overall effect: Z = 12.62 (P < 0.00001)02 OCD-CPRS Thoren 1980a 6.38(2.75) 8.75(3.18) -0.75 [-1.78, 0.27] Subtotal (95% CD) 8 -0.75 [-1.78, 0.27] Test for heterogeneity; not applicable Test for overall effect: Z = 1.44 (P = 0.15) 03 6-item OCD Scale - amelioration -1.70 [-2.99, -0.42] -1.70 [-2.99, -0.42] Montgomery 1990 -10.00(5.11) -0.71(5.11)Subtotal (95% CI) Test for heterogeneity: not applicable Test for overall effect: Z = 2.60 (P = 0.009)438 -0.93 [-1.07, -0.79] Test for heterogeneity: Chi² = 49.22, df = 5 (P < 0.00001), l² = 89.8% Test for overall effect: Z = 12.91 (P < 0.00001)

-2

Ó Favours Clomipramine Favours Placebo

Removing heterogeneity from Clomipramine v Placebo (Y-BOCS/CPRS): Burnham1993 and Zohar1996a removed because they reported change scores



Favours Clomipramine

Favours placebo

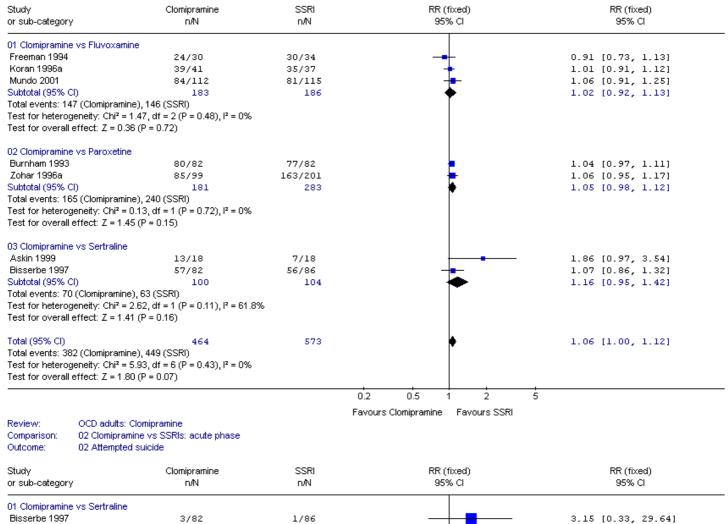
Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo: acute phase Outcome 16 Leyton Obsessional Inventory: resistance SMD (fixed) SMD (fixed) Study Placebo or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI Thoren 1980a 8 -13.00(19.60) 8 -4.30(18.60) -0.43 [-1.43, 0.56] Total (95% CI) -0.43 [-1.43, 0.56] Test for heterogeneity: not applicable Test for overall effect: Z = 0.85 (P = 0.40) -10 -5 ò 10 Favours Clomipramine Favours placebo OCD adults: Clomipramine Review: Comparison: 01 Clomipramine vs placebo: acute phase Outcome 17 Leyton Obsessional Inventory: interference SMD (fixed) SMD (fixed) Placebo Study Clomipramine Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Thoren 1980a 8 -12.30(24.00) -8.40(15.90) -0.18 [-1.16, 0.80] Total (95% CI) -0.18 [-1.16, 0.80] Test for heterogeneity: not applicable Test for overall effect: Z = 0.36 (P = 0.72) -10 -5 10 0 Favours Clomipramine Favours placebo Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo; acute phase Outcome 18 Self-Rating Obsessive-Compulsive Personality Inventory Study Clomipramine Placebo SMD (fixed) SMD (fixed) Ν Ν or sub-category Mean (SD) Mean (SD) 95% CI 95% CI Stein 1992 21 45.17(7.36) 23 49.31(7.36) -0.55 [-1.16, 0.05] Total (95% CI) 21 23 -0.55 [-1.16, 0.05] Test for heterogeneity; not applicable Test for overall effect: Z = 1.79 (P = 0.07) -10 -5 ń 10 Favours Clomipramine Favours Placebo Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo: acute phase Outcome 19 Montgomery-Asberg Depression Rating Scale Study Clomipramine Placebo SMD (fixed) SMD (fixed) or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI Zohar 1996a 94 -2.40(7.30) 99 -0.60(7.90) -0.24 [-0.52, 0.05] Total (95% CI) -0.24 [-0.52, 0.05] Test for heterogeneity: not applicable Test for overall effect: Z = 1.63 (P = 0.10) 10 -10 Ó Favours Clomipramine Favours Placebo OCD adults: Clomipramine Review: 01 Clomipramine vs placebo: acute phase Comparison: Outcome 20 Clinical Global Impressions: severity of illness SMD (fixed) SMD (fixed) Study Clomipramine Placebo Ν Mean (SD) 95% CI or sub-category Ν Mean (SD) 95% CI Zohar 1996a -1.10(1.20) 99 -0.70(1.30) -0.32 [-0.60, -0.03] Total (95% CI) 99 -0.32 [-0.60, -0.03] Test for heterogeneity: not applicable Test for overall effect: Z = 2.20 (P = 0.03) -10 10 -5 0 Favours Clomipramine Favours Placebo Review: OCD adults: Clomipramine Comparison: 01 Clomipramine vs placebo: acute phase Outcome 21 Symptom Checklist-90 Placebo SMD (fixed) SMD (fixed) Study or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI Zohar 1996a -31.90(39.90) 77 -19.50(37.30) -0.32 [-0.64, 0.00] Total (95% CI) 77 -0.32 [-0.64, 0.00] Test for heterogeneity: not applicable Test for overall effect: Z = 1.95 (P = 0.05) -10 -5 Ó 5 10

Favours Clomipramine Favours Placebo

Review: OCD adults: Clomipramine
Comparison: 02 Clomipramine vs SSRIs: acute phase
Outcome: 01 Adverse events

Subtotal (95% CI)

Total events: 3 (Clomipramine), 1 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 1.00 (P = 0.32)



86

0.01

0.1

Favours Clomipramine

3.15 [0.33, 29.64]

100

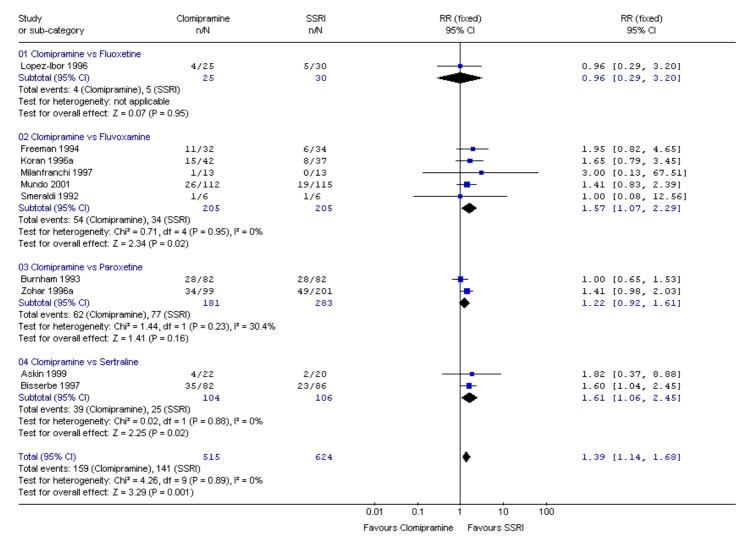
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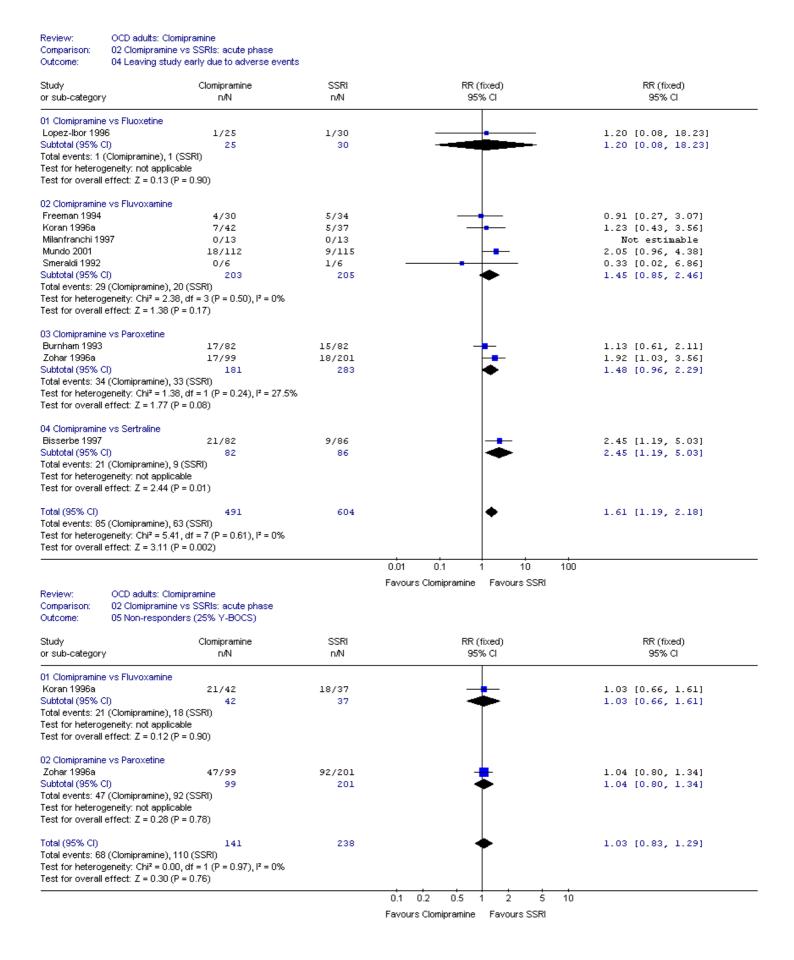
Favours SSRI

Review: OCD adults: Clomipramine

Comparison: 02 Clomipramine vs SSRIs: acute phase

Outcome: 03 Leaving study early



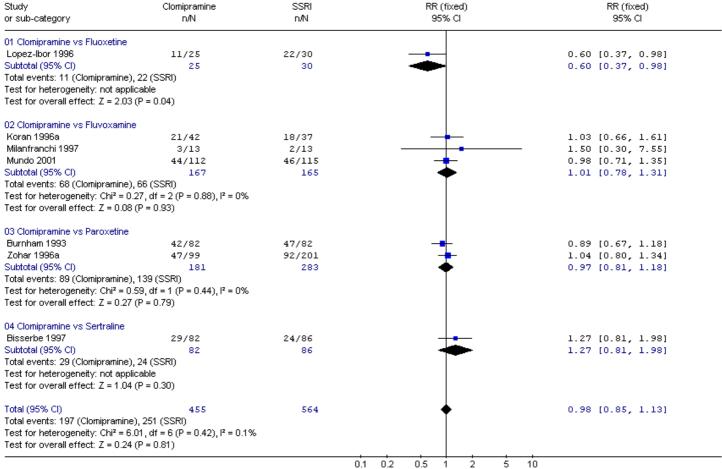


Review: OCD adults: Clomipramine 02 Clomipramine vs SSRIs: acute phase Comparison: Outcome: 06 Non-responders (35% Y-BOCS) SSRI RR (fixed) RR (fixed) Study Clomipramine or sub-category n/N n/N 95% CI 95% CI 01 Clomipramine vs Fluvoxamine Mundo 2001 44/112 46/115 0.98 [0.71, 1.35] Subtotal (95% CI) 112 115 0.98 [0.71, 1.35] Total events: 44 (Clomipramine), 46 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.11 (P = 0.91) 0.1 0.2 0.5 2 5 10 Favours Clomipramine Favours SSRI Review: OCD adults: Clomipramine Comparison: 02 Clomipramine vs SSRIs: acute phase Outcome: 07 Non-responders (CGI; 25% Y-BOCS) Study Clomipramine SSRI RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 Clomipramine vs Fluoxetine Lopez-Ibor 1996 11/25 22/30 0.60 [0.37, 0.98] 0.60 [0.37, 0.98] Subtotal (95% CI) 25 30 Total events: 11 (Clomipramine), 22 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 2.03 (P = 0.04) 02 Clomipramine vs Paroxetine 0.89 [0.67, 1.18] Burnham 1993 42/82 47/82 Subtotal (95% CD 0.89 [0.67, 1.18] 82 82 Total events: 42 (Clomipramine), 47 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.78 (P = 0.43) Total (95% CI) 0.81 [0.63, 1.03] 112 107 Total events: 53 (Clomipramine), 69 (SSRI) Test for heterogeneity: $Chi^2 = 1.90$, df = 1 (P = 0.17), $I^2 = 47.3\%$ Test for overall effect: Z = 1.73 (P = 0.08) 0.2 0.5 ż 5 10 Favours Clomipramine Favours SSRI OCD adults: Clomipramine Review: 02 Clomipramine vs SSRIs; acute phase Comparison: Outcome: 09 Non-responders (CGI-I) SSRI RR (fixed) RR (fixed) Study Clomipramine 95% CI or sub-category n/N n/N 95% CI 01 Clomipramine vs Fluvoxamine Milanfranchi 1997 3/13 2/13 1.50 [0.30, 7.55] Subtotal (95% CI) 1.50 [0.30, 7.55] 13 Total events: 3 (Clomipramine), 2 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.49 (P = 0.62) 02 Clomipramine vs Sertraline Bisserbe 1997 29/82 24/86 1.27 [0.81, 1.98] Subtotal (95% CI) 86 1.27 [0.81, 1.98] Total events: 29 (Clomipramine), 24 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 1.04 (P = 0.30) Total (95% CI) 99 1.29 [0.83, 1.98] Total events: 32 (Clomipramine), 26 (SSRI) Test for heterogeneity: $Chi^2 = 0.04$, df = 1 (P = 0.84), $I^2 = 0\%$ Test for overall effect: Z = 1.14 (P = 0.25) 0.2 10 0.5 5

Favours Clomipramine

Favours SSRI

Review: OCD adults: Clomipramine
Comparison: 02 Clomipramine vs SSRIs: acute phase
0utcome: 10 Non-responders (all)
Study Clomipramine



Review: OCD adults: Clomipramine

Comparison: 02 Clomipramine vs SSRIs; acute phase

Outcome: 11 Y-BOCS

Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Clomipramine vs Fluoxetine						
Lopez-lbor 1996	24	-8.90(7.13)	30	-7.50(9.29)		-0.16 [-0.70, 0.37]
Subtotal (95% CI)	24		30			-0.16 [-0.70, 0.37]
est for heterogeneity: not app	licable					
Test for overall effect: $Z = 0.60$	(P = 0.55)					
02 Clomipramine vs Fluvoxamin	ie					
Koran 1996a	39	17.00(8.55)	34	17.80(7.70)		-0.10 [-0.56, 0.36]
Milanfranchi 1997	12	16.50(11.00)	13	18.40(9.20)		-0.18 [-0.97, 0.60]
Mundo 2001	61	12.50(8.00)	67	14.70(10.10)		-0.24 [-0.59, 0.11]
Smeraldi 1992	5	15.20(12.40)	5	18.20(2.50)	-	-0.30 [-1.55, 0.95]
Subtotal (95% CI)	117		119			-0.19 [-0.45, 0.06]
[est for heterogeneity: Chi² = 0 [est for overall effect: Z = 1.48		P = 0.97), I ² = 0%				
3 Clomipramine vs Paroxetine						
Burnham 1993	78	-7.73(7.42)	79	-5.61(7.47)		-0.28 [-0.60, 0.03]
Zohar 1996a	94	-8.00(8.20)	198	-8.00(8.00)		0.00 [-0.25, 0.25]
Subtotal (95% CI)	172		277		-	-0.11 [-0.30, 0.09]
[est for heterogeneity: Chi² = 1 [est for overall effect: Z = 1.09		P = 0.16), I ² = 48.4%				
Total (95% CI)	313		426		•	-0.14 [-0.29, 0.01]
[est for heterogeneity: Chi² = 2 [est for overall effect: Z = 1.85		° = 0.87), l² = 0%				
					-1 -0.5 0 0.5	1
					Favours Clomipramine Favours S	CPI

Favours Clomipramine

Favours SSRI

Review: OCD adults: Clomipramine

02 Clomipramine vs SSRIs: acute phase 12 NIMH-OC Comparison:

Outcome:

lomipramine Mean (SD)	N	SSRI Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
5.90(2.80)	112	5.90(3.10)		0.00 [-0.27, 0.27]
	112		-	0.00 [-0.27, 0.27]
			T	
-2.06(2.03)	79	-1.42(2.04)		-0.31 [-0.63, 0.00]
-2.50(2.60)	198	-2.50(2.90)		0.00 [-0.25, 0.25]
	277		-	-0.12 [-0.31, 0.08]
0.12), I² = 57.6%			-	
	389		•	-0.08 [-0.23, 0.08]
0.24), I ² = 30.0%			٦	
• •				
0.24), l² =	30.0%			

Favours Clomipramine Favours SSRI

Review:

OCD adults: Clomipramine 02 Clomipramine vs SSRIs: acute phase Comparison:

Outcome: 13 Comprehensive Psychopathological Rating Scale: obsessive-compulsive

Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)			MD (fixed 95% CI)		SMD (fixed) 95% Cl
01 Clomipramine vs Fluoxet	ine									
Lopez-lbor 1996	24	-10.90(7.33)	30	-8.10(7.64)		_	-			-0.37 [-0.91, 0.17]
Subtotal (95% CI)	24		30							-0.37 [-0.91, 0.17]
Test for heterogeneity: not	applicable									
Test for overall effect: Z =	1.33 (P = 0.18)									
					- 1	-0.5	_	0.5	+	
					-1	-0.5	U	0.5	1	

Favours Clomipramine Favours SSRI

Review: OCD adults: Clomipramine

Comparison: 02 Clomipramine vs SSRIs: acute phase Outcome: 14 Hamilton Rating Scale for Depression

Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Clomipramine vs Fluoxetino	e					
Lopez-lbor 1996	24	-9.40(7.71)	30	-7.50(5.93)		-0.28 [-0.82, 0.26]
Subtotal (95% CI)	24		30			-0.28 [-0.82, 0.26]
Test for heterogeneity: not as	oplicable					
Test for overall effect: Z = 1.	00 (P = 0.32)					
02 Clomipramine vs Fluvoxam	nine					
Koran 1996a	39	5.80(4.70)	34	6.10(3.30)		-0.07 [-0.53, 0.39]
Mundo 2001	105	7.00(4.50)	112	7.70(6.00)		-0.13 [-0.40, 0.14]
Subtotal (95% CI)	144		146			-0.12 [-0.35, 0.11]
Test for heterogeneity: Chi2 =	0.05, df = 1 (F	P = 0.83), I ² = 0%			-	
Test for overall effect: $Z = 0$.	99 (P = 0.32)					
Total (95% CI)	168		176			-0.14 [-0.35, 0.07]
Test for heterogeneity: Chi2 =	0.33, df = 2 (F	P = 0.85), I ² = 0%			_	
Test for overall effect: Z = 1.3	30 (P = 0.19)	**				
					1 -0.5 0 0.5	<u> </u>

Review: OCD adults: Clomipramine Comparison: 02 Clomipramine vs SSRIs: acute phase Outcome 15 Montgomery-Asberg Depression Rating Scale SMD (fixed) SMD (fixed) Study SSRI or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI 01 Clomipramine vs Fluoxetine Lopez-lbor 1996 24 -13.80(9.09) 30 -10.10(8.20) -0.42 [-0.97, 0.12] Subtotal (95% CI) 24 30 -0.42 [-0.97, 0.12] Test for heterogeneity: not applicable Test for overall effect: Z = 1.53 (P = 0.13) 02 Clomipramine vs Fluvoxamina Smeraldi 1992 5 9.60(10.30) 5 11.80(6.80) -0.23 [-1.47, 1.02] Subtotal (95% CI) -0.23 [-1.47, 1.02] 5 Test for heterogeneity: not applicable Test for overall effect: Z = 0.36 (P = 0.72) 03 Clomipramine vs Paroxetine Zohar 1996a 94 -2.40(7.30) 198 -3.40(7.90) 0.13 [-0.12, 0.38] Subtotal (95% CI) 0.13 [-0.12, 0.38] Test for heterogeneity: not applicable Test for overall effect: Z = 1.03 (P = 0.30) 233 0.03 [-0.19, 0.25] Test for heterogeneity: $Chi^2 = 3.47$, df = 2 (P = 0.18), $I^2 = 42.4\%$ Test for overall effect: Z = 0.24 (P = 0.81) Favours Clomipramine Favours SSRI Review: OCD adults: Clomipramine 02 Clomipramine vs SSRIs: acute phase Comparison: Outcome 16 Depression (HRSD; MADRS) SSRI SMD (fixed) SMD (fixed) Study Clomipramine Mean (SD) N Mean (SD) 95% CI 95% CI or sub-category Ν 01 Clomipramine vs Fluvoxamine Koran 1996a 39 5.80(4.70) 34 6.10(3.30) -0.07 [-0.53, 0.39] Mundo 2001 -0.13 [-0.40, 0.14] -0.23 [-1.47, 1.02] 105 7.00(4.50) 112 7.70(6.00) Smeraldi 1992 9.60(10.30) 11.80(6.80) 5 Subtotal (95% CI) 149 151 -0.12 [-0.35, 0.11] Test for heterogeneity: $Chi^2 = 0.08$, df = 2 (P = 0.96), $I^2 = 0\%$ Test for overall effect: Z = 1.04 (P = 0.30) 02 Clomipramine vs Paroxetine Zohar 1996a 94 -2.40(7.30) 198 -3.40(7.90)0.13 [-0.12, 0.38] Subtotal (95% CI) 0.13 [-0.12, 0.38] 94 198 Test for heterogeneity: not applicable Test for overall effect: Z = 1.03 (P = 0.30) Total (95% CI) 349 -0.01 [-0.17, 0.16] Test for heterogeneity: Chi² = 2.21, df = 3 (P = 0.53), l² = 0% Test for overall effect: Z = 0.06 (P = 0.95) -2 Ò Favours Clomipramine Favours SSRI OCD adults: Clomipramine Review: Comparison: 02 Clomipramine vs SSRIs; acute phase Outcome 17 Clinical Anxiety Scale SSRI SMD (fixed) SMD (fixed) or sub-category M Mean (SD) M Mean (SD) 95% CI 95% CI 01 Clomipramine vs Fluvoxamine Mundo 2001 -0.07 [-0.33, 0.20] 6.70(4.30) 7.00(4.60) Subtotal (95% CI) 105 112 -0.07 [-0.33, 0.20] Test for heterogeneity; not applicable Test for overall effect; Z = 0.49 (P = 0.62) -0.5 Ó 0.5 Favours Clomipramine Favours SSRI Review: OCD adults: Clomipramine 02 Clomipramine vs SSRIs; acute phase Comparison: Outcome 18 Covi Anxiety Scale Study SSRI SMD (fixed) SMD (fixed) Clomipramine or sub-category Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Clomipramine vs Fluoxetine 0.00 [-0.54, 0.54] Lopez-Ibor 1996 24 -2.40(3.09) 30 -2.40(3.10) 0.00 [-0.54, 0.54] Subtotal (95% CI) 30 Test for heterogeneity: not applicable Test for overall effect: Z = 0.00 (P = 1.00) 10 Ó Favours Clomipramine Favours SSRI

Review: OCD adults: Clomipramine

Comparison: 02 Clomipramine vs SSRIs; acute phase

Outcome: 19 Anxiety (Clinical Anxiety Scale; Covi Anxiety Scale)

Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Clomipramine vs Fluvox	camine					
Mundo 2001	105	6.70(4.30)	112	7.00(4.60)		-0.07 [-0.33, 0.20]
Subtotal (95% CI)	105		112			-0.07 [-0.33, 0.20]
Test for heterogeneity: not	t applicable				٦	
Test for overall effect: Z =	0.49 (P = 0.62)					
02 Clomipramine vs Fluoxe	etine					
Lopez-lbor 1996	24	-2.40(3.09)	30	-2.40(3.10)		0.00 [-0.54, 0.54]
Subtotal (95% CI)	24		30			0.00 [-0.54, 0.54]
Test for heterogeneity: not	t applicable				T	
Test for overall effect: Z =	0.00 (P = 1.00)					
Total (95% CI)	129		142			-0.05 [-0.29, 0.18]
Test for heterogeneity: Chi	$i^2 = 0.05$, $df = 1$ (1)	P = 0.83), I ² = 0%				
Test for overall effect: Z =	0.44 (P = 0.66)					
					-1 -0.5 0 0.5	
					-1 -0.5 0 0.5	ı

Favours Clomipramine Favours SSRI

Review: OCD adults: Clomipramine

Comparison: 02 Clomipramine vs SSRIs: acute phase 20 Clinical Global Impressions: global improvement Outcome:

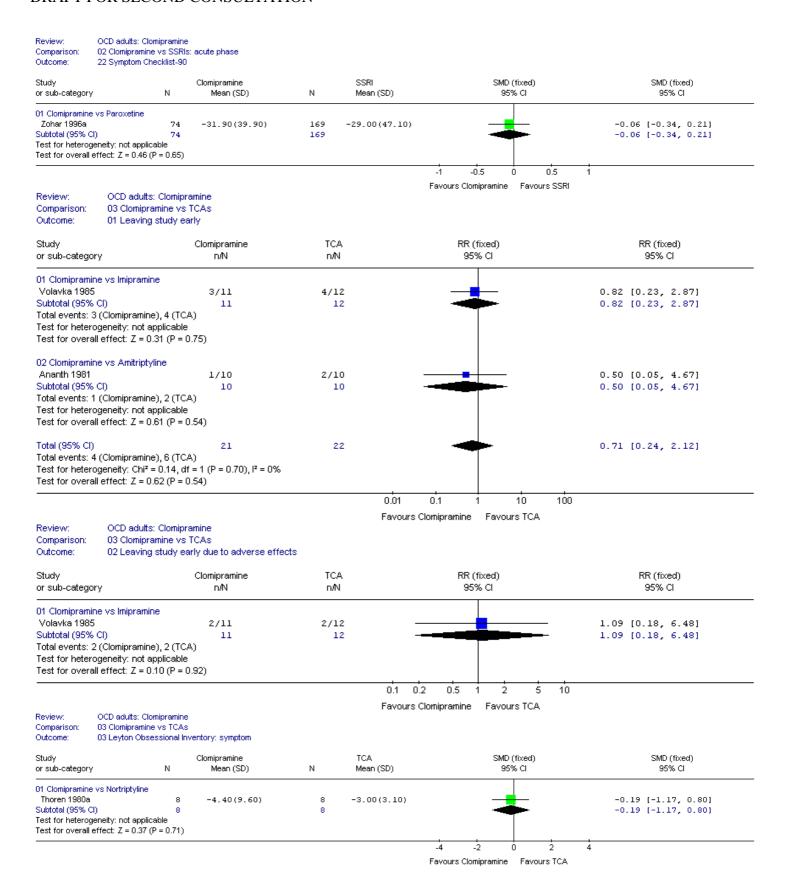
Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Clomipramine vs Fluoxetin	B					
Lopez-lbor 1996	24	-2.60(1.19)	30	-2.60(1.13)		0.00 [-0.54, 0.54]
Subtotal (95% CI)	24		30			0.00 [-0.54, 0.54]
Test for heterogeneity: not ap	plicable				T	
Test for overall effect: $Z = 0$.	00 (P = 1.00)					
02 Clomipramine vs Fluvoxam	ine					
Mundo 2001	105	2.30(1.00)	112	2.30(1.00)		0.00 [-0.27, 0.27]
Subtotal (95% CI)	105		112			0.00 [-0.27, 0.27]
Test for heterogeneity: not ap	plicable				Ī	
Test for overall effect: $Z = 0$.	00 (P = 1.00)					
Total (95% CI)	129		142		•	0.00 [-0.24, 0.24]
Test for heterogeneity: Chi² =	0.00, df = 1	P = 1.00), I ² = 0%			T	
Test for overall effect: $Z = 0.1$	00 (P = 1.00)					

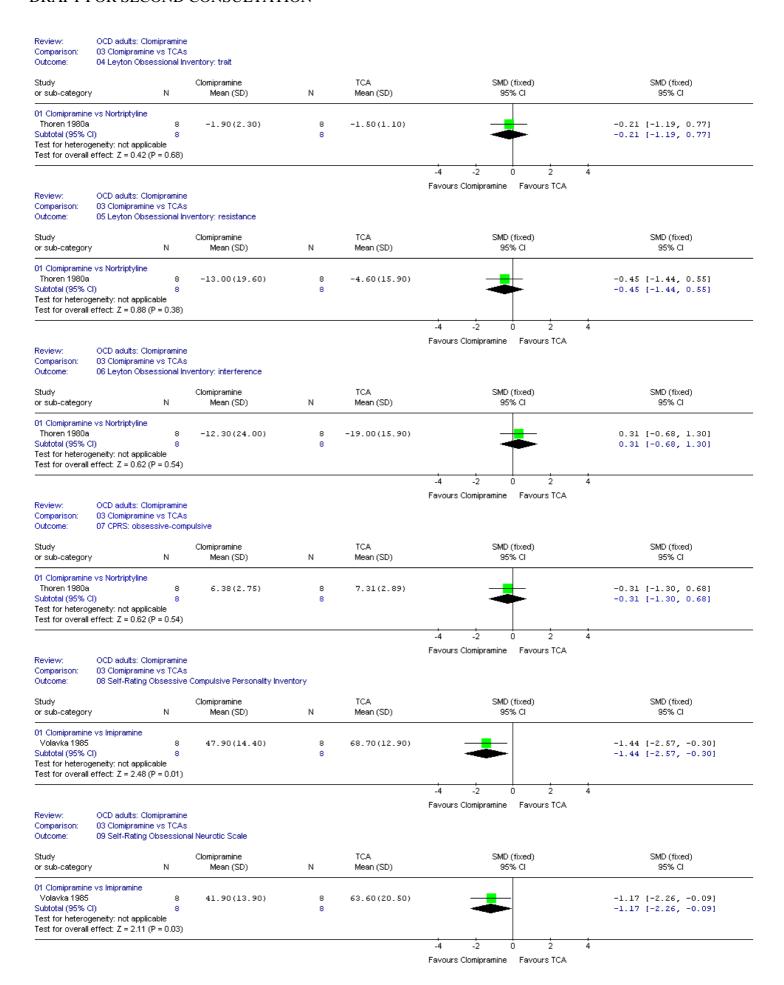
Favours Clomipramine Favours SSRI Review: OCD adults: Clomipramine

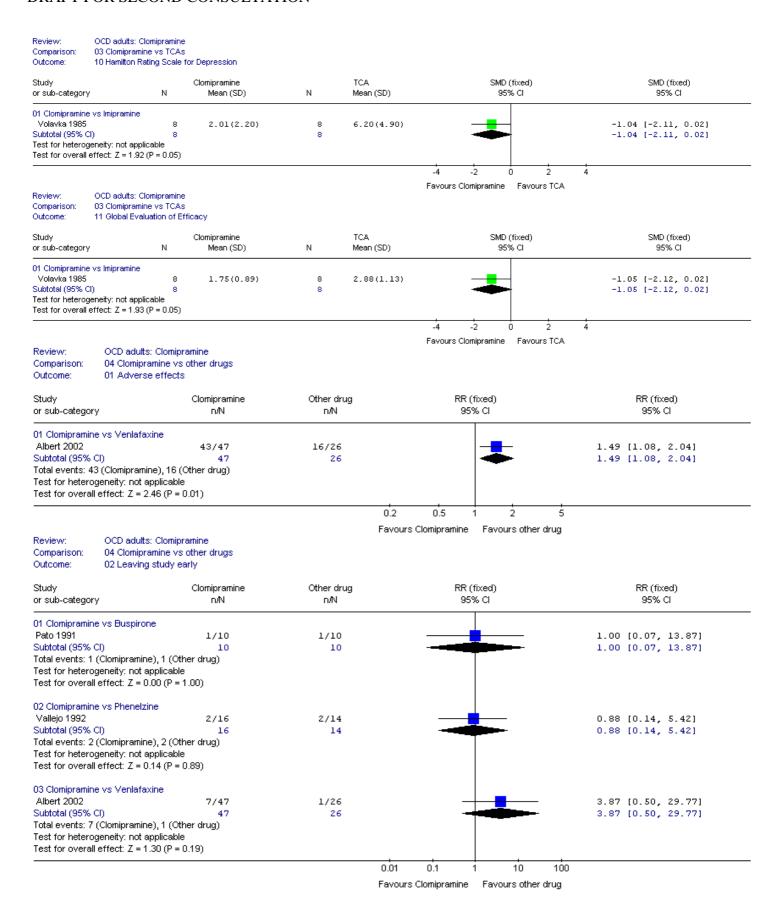
02 Clomipramine vs SSRIs: acute phase 21 Clinical Global Impressions: severity of illness Comparison:

Outcome:

Study or sub-category	N	Clomipramine Mean (SD)	N	SSRI Mean (SD)			ID (fixed) 95% Cl			SMD (fixed) 95% Cl
01 Clomipramine vs Fluoxetine										
Lopez-Ibor 1996	24	-1.90(1.14)	30	-0.90(1.23)			-			-0.83 [-1.39, -0.27]
Subtotal (95% CI)	24		30			•	▶			-0.83 [-1.39, -0.27]
Test for heterogeneity: not app	olicable									
Test for overall effect: $Z = 2.8$	9 (P = 0.004)									
02 Clomipramine vs Fluvoxami	ne									
Mundo 2001	105	3.20(1.20)	112	3.40(1.30)			=			-0.16 [-0.43, 0.11]
Subtotal (95% CI)	105		112				*			-0.16 [-0.43, 0.11]
Test for heterogeneity: not app	olicable									
Test for overall effect: $Z = 1.1$	7 (P = 0.24)									
03 Clomipramine vs Paroxetine	!									
Zohar 1996a	94	-1.10(1.20)	194	-1.20(1.40)			+			0.07 [-0.17, 0.32]
Subtotal (95% CI)	94		194				•			0.07 [-0.17, 0.32]
Test for heterogeneity: not app	olicable									
Test for overall effect: $Z = 0.5$	9 (P = 0.55)									
Fotal (95% CI)	223		336				•			-0.11 [-0.28, 0.06]
Test for heterogeneity: Chi² = :	3.57, $df = 2$ (1	P = 0.01), I ² = 76.7%					1			•
Test for overall effect: $Z = 1.2$										
					-4	-2	0	2	4	
					Favours (Clomipramin	e Favo	urs SSRI		

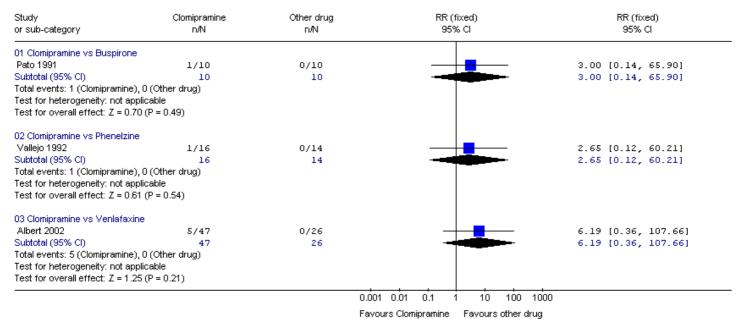






Review: OCD adults: Clomipramine Comparison: 04 Clomipramine vs other drugs

Outcome: 03 Leaving study early due to adverse effects



Review: OCD adults: Clomipramine Comparison: 04 Clomipramine vs other drugs

Outcome: 04 Non-responders

Study or sub-category	Clomipramine n/N	Other drug n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Clomipramine vs Venlafa	axine			
Albert 2002	27/47	17/26		0.88 [0.61, 1.28]
Subtotal (95% CI)	47	26		0.88 [0.61, 1.28]
Total events: 27 (Clomipram	ine), 17 (Other drug)			
Test for heterogeneity: not	applicable			
Test for overall effect: Z = I	0.68 (P = 0.50)			
			0.5 0.7 1 1.5	2

OCD adults: Clomipramine Favours other drug

Review: OCD adults: Clomipramine Comparison: 04 Clomipramine vs other drugs

Outcome: 05 Y-BOCS

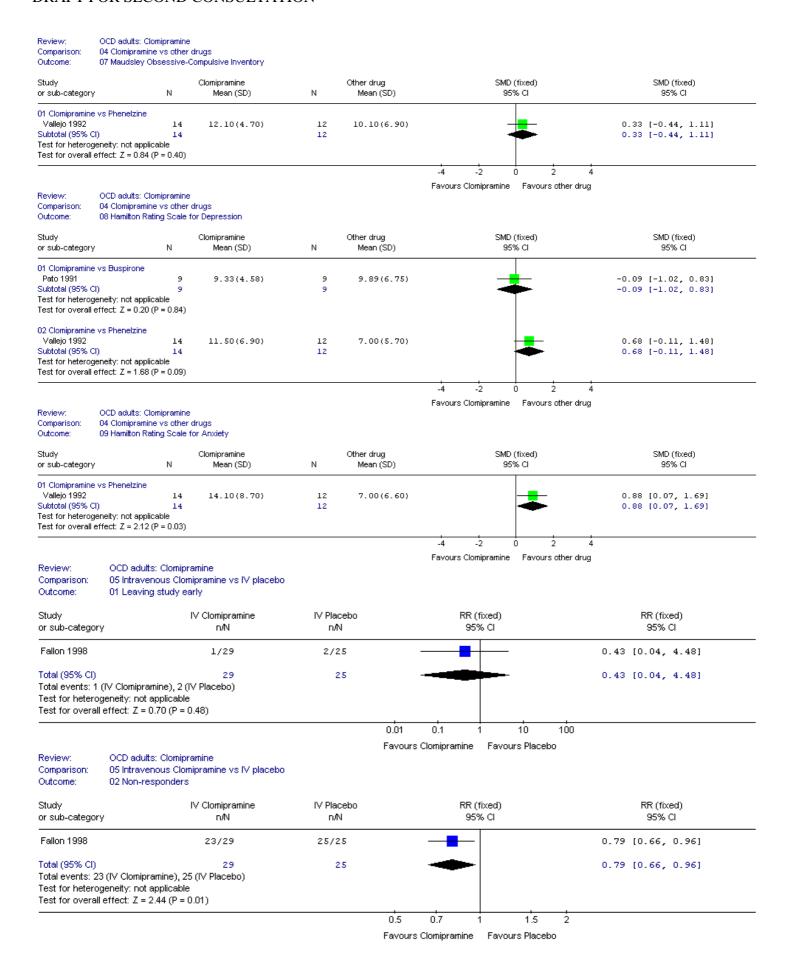
Study or sub-category	N	Clomipramine Mean (SD)	N	Other drug Mean (SD)		SMD (fixed) 95% Cl	SMD (fixed) 95% CI
01 Clomipramine vs Buspiron	ie						
Pato 1991	9	17.67(4.97)	9	16.78(6.46)		_	0.15 [-0.78, 1.07]
Subtotal (95% CI)	9		9				0.15 [-0.78, 1.07]
Test for heterogeneity: not a	pplicable					_	•
Test for overall effect: $Z = 0$.	31 (P = 0.76)						
02 Clomipramine vs Venlafax	tine						
Albert 2002	47	17.30(6.15)	26	18.36(7.11)		-	-0.16 [-0.64, 0.32]
Subtotal (95% CI)	47		26			•	-0.16 [-0.64, 0.32]
Test for heterogeneity: not a	pplicable					1	·
Test for overall effect: $Z = 0$.	.66 (P = 0.51)						
					-4 -2) 0 2	4

Review: OCD adults: Clomipramine Favours other drug

Comparison: 04 Clomipramine vs other drugs

Outcome: 06 NIMH-OC

Study or sub-category	N	Clomipramine Mean (SD)	N	Other drug Mean (SD)		:	SMD (fixed 95% CI)		SMD (fixed) 95% Cl
01 Clomipramine vs Buspiror	ne									
Pato 1991	9	5.22(1.20)	9	5.11(1.27)			-			0.08 [-0.84, 1.01]
Subtotal (95% CI)	9		9				-			0.08 [-0.84, 1.01]
est for heterogeneity: not a	applicable						Ī			
「est for overall effect: Z = 0	0.18 (P = 0.86)									
					-4	-2	6	2	4	
					Favours	Clomiprar	nine Fav	ours other	r drug	



OCD adults: Clomipramine Review: Comparison: 05 Intravenous Clomipramine vs IV placebo Outcome 03 Y-BOCS IV Placebo SMD (fixed) SMD (fixed) Study IV Clomipramine or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI Fallon 1998 28 25,20(7,00) 23 26.10(5.00) -0.14 [-0.70. 0.41] Total (95% CI) 23 -0.14 [-0.70, 0.41] Test for heterogeneity: not applicable Test for overall effect: Z = 0.51 (P = 0.61) -0.5 ò 0.5 Favours Clomipramine Favours Placebo OCD adults: Clomipramine Review: Comparison: 05 Intravenous Clomipramine vs IV placebo Outcome 04 NIMH-OC IV Placebo SMD (fixed) SMD (fixed) Study IV Clomipramine N Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Fallon 1998 28 10.40(2.00) 23 10.80(1.00) -0.24 [-0.80, 0.31] Total (95% CI) 23 -0.24 [-0.80, 0.31] 28 Test for heterogeneity: not applicable Test for overall effect: Z = 0.86 (P = 0.39) -0.5 0.5 Ó Favours Clomipramine Favours Placebo Review: OCD adults: Clomipramine 05 Intravenous Clomipramine vs IV placebo Comparison: Outcome: 05 Clinical Global Impressions: severity Study IV Clomipramine IV Placebo SMD (fixed) SMD (fixed) Ν Ν Mean (SD) or sub-category Mean (SD) 95% CI 95% CI Fallon 1998 28 5.30(1.00) 23 5.70(5.00) -0.11 [-0.67, 0.44] Total (95% CI) 28 23 -0.11 [-0.67, 0.44] Test for heterogeneity; not applicable Test for overall effect: Z = 0.41 (P = 0.68) -0.5 ń 0.5 Favours Clomipramine Favours Placebo OCD adults: Clomipramine Review: Comparison: 05 Intravenous Clomipramine vs IV placebo Outcome 06 Hamilton Rating Scale for Depression IV Placebo SMD (fixed) SMD (fixed) Study IV Clomipramine or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI Fallon 1998 12.70(7.00) 28 12.00(6.00) 23 -0.11 [-0.66, 0.45] Total (95% CI) 23 -0.11 [-0.66, 0.45] Test for heterogeneity: not applicable Test for overall effect: Z = 0.38 (P = 0.71) -0.5 0.5 Favours Clomipramine Favours Placebo Review: OCD adults: Clomipramine Comparison: 06 Intravenous Clomipramine vs oral Clomipramine Outcome: 01 Adverse effects Oral Clomipramine RR (fixed) RR (fixed) Study IV Clomipramine 95% CI 95% CI or sub-category n/N n/N Koran 1997 4/7 5/8 0.91 [0.40, 2.11]

8

0.2

0.5

Favours IV

0.91 [0.40, 2.11]

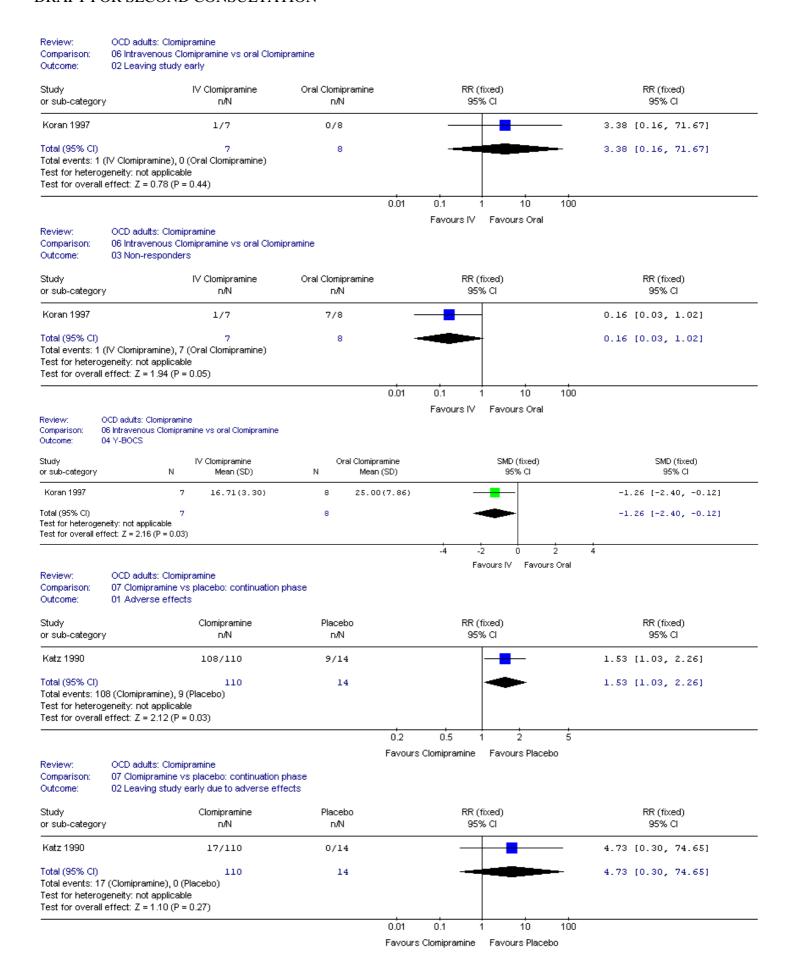
5

Favours Oral

Total (95% CI)

Total events: 4 (IV Clomipramine), 5 (Oral Clomipramine)

Test for heterogeneity: not applicable Test for overall effect: Z = 0.21 (P = 0.83)



Review: OCD adults: Clomipramine Comparison: 07 Clomipramine vs placebo: continuation phase Outcome 03 NIMH-OC SMD (fixed) SMD (fixed) Study Clomipramine Placebo or sub-category N Mean (SD) N Mean (SD) 95% CI 95% CI Katz 1990 101 5.10(2.70) 12 9.00(2.50) -1.44 [-2.07. -0.82] Total (95% CI) 12 -1.44 [-2.07, -0.82] Test for heterogeneity: not applicable Test for overall effect: Z = 4.51 (P < 0.00001) ò Favours Clomipramine Favours Placebo OCD adults: Clomipramine Review: Comparison: 07 Clomipramine vs placebo: continuation phase Outcome 04 Physician's Global Evaluation SMD (fixed) Study Placebo SMD (fixed) Clomipramine Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category Katz 1990 101 1.89(0.94) 12 3.33(0.89) -1.53 [-2.16, -0.90] Total (95% CI) 12 -1.53 [-2.16, -0.90] 101 Test for heterogeneity: not applicable Test for overall effect: Z = 4.74 (P < 0.00001)Ó Favours Clomipramine Favours Placebo OCD adults: Clomipramine Comparison: 08 Clomipramine vs SSRIs: continuation phase 01 Leaving study early Outcome: Study Clomipramine SSRI RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 Clomipramine 200mg vs Fluoxetine 60mg (non-responders) 0.88 [0.30, 2.58] Lopez-lbor 1996 6/14 3/8 Subtotal (95% CI) 14 0.88 [0.30, 2.58] Total events: 3 (Clomipramine), 6 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.24 (P = 0.81) 02 Clomipramine 100mg vs Fluoxetine 20mg (responders) 1.69 [0.55, 5.24] Lopez-lbor 1996 3/11 6/13 Subtotal (95% CI) 1.3 1.1 1.69 [0.55, 5.24] Total events: 6 (Clomipramine), 3 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.91 (P = 0.36) 0.2 0.5 5 10 Favours Clomipramine Favours SSRI Review: OCD adults: Clomipramine Comparison: 08 Clomipramine vs SSRIs: continuation phase Outcome: 02 Leaving study early due to adverse effects SSRI Study RR (fixed) RR (fixed) Clomipramine or sub-category n/Ν n/Ν 95% CL 95% CL 01 Clomipramine 200mg vs Fluoxetine 60mg (non-responders) Lopez-Ibor 1996 0/8 2/14 0.33 [0.02, 6.19] Subtotal (95% CI) 0.33 [0.02, 6.19] 14 Total events: 0 (Clomipramine), 2 (SSRI) Test for heterogeneity; not applicable Test for overall effect: Z = 0.74 (P = 0.46) 02 Clomipramine 100mg vs Fluoxetine 20mg (responders) Lopez-lbor 1996 1/11 0.85 [0.06, 12.01] 1/13 Subtotal (95% CI) 13 11 0.85 [0.06, 12.01] Total events: 1 (Clomipramine), 1 (SSRI) Test for heterogeneity: not applicable Test for overall effect: Z = 0.12 (P = 0.90)

0.01

0.1

Favours Clomipramine

10

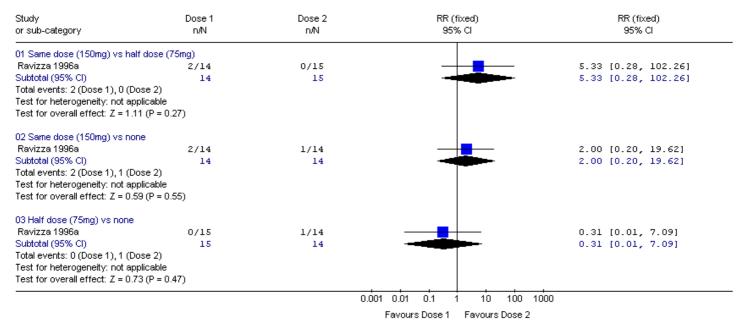
Favours SSRI

100

Review: OCD adults: Clomipramine

Comparison: 09 Clomipramine same dose vs half dose vs none: continuation/discontinuation

Outcome: 01 Leaving study early



Review: OCD adults: Clomipramine

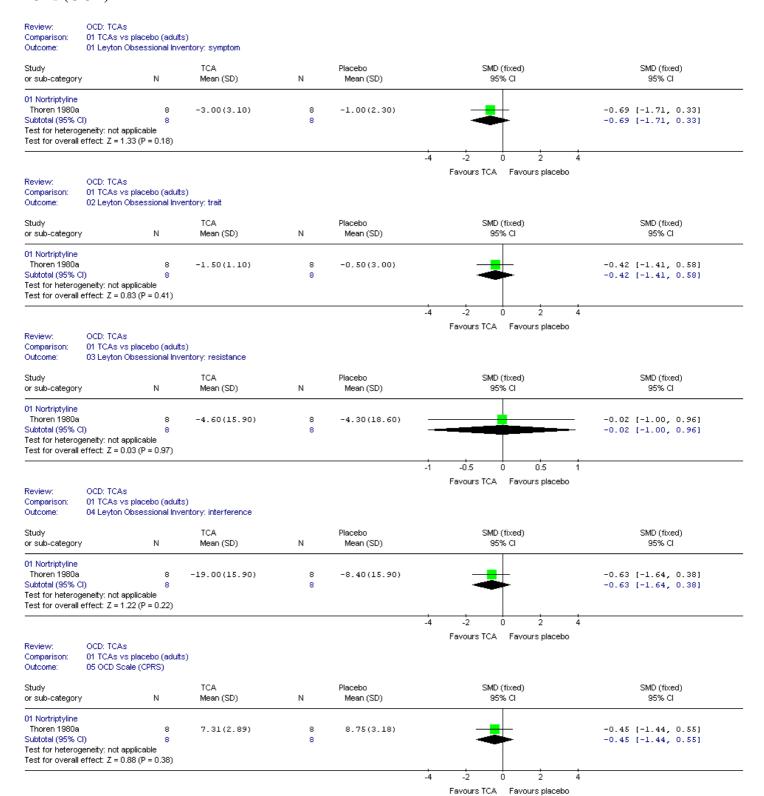
Comparison: 09 Clomipramine same dose vs half dose vs none: continuation/discontinuation

Outcome: 02 Relapse (CGI; 25% Y-BOCS)

Study or sub-category	Dose 1 n/N	Dose 2 n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Same dose (150mg) vs half de	ose (75ma)			
Ravizza 1996a	5/14	4/15		1.34 [0.45, 4.00]
Subtotal (95% CI)	14	15		1.34 [0.45, 4.00]
rotal events: 5 (Dose 1), 4 (Dose	2)			
est for heterogeneity; not applic	,			
est for overall effect: Z = 0.52 (P = 0.60)			
02 Same dose (150mg) vs none				
Ravizza 1996a	5/14	11/14		0.45 [0.21, 0.97]
Subtotal (95% CI)	14	14		0.45 [0.21, 0.97]
otal events: 5 (Dose 1), 11 (Dos	se 2)			
est for heterogeneity: not applic	able			
est for overall effect: Z = 2.05 (P = 0.04)			
3 Half dose (75mg) vs none			_	
Ravizza 1996a	4/15	11/14		0.34 [0.14, 0.82]
Subtotal (95% CI)	15	14		0.34 [0.14, 0.82]
'otal events: 4 (Dose 1), 11 (Dos	,			
'est for heterogeneity: not applic				
est for overall effect: Z = 2.40 (P = 0.02)			
04 Continuation v discontinuation			_	
Ravizza 1996a	9/29	11/14		0.39 [0.22, 0.73]
Subtotal (95% CI)	29	14		0.39 [0.22, 0.73]
'otal events: 9 (Dose 1), 11 (Dos	,			
est for heterogeneity: not applic				
est for overall effect: Z = 3.00 (P = 0.003)			

Favours Dose 1 Favours Dose 2

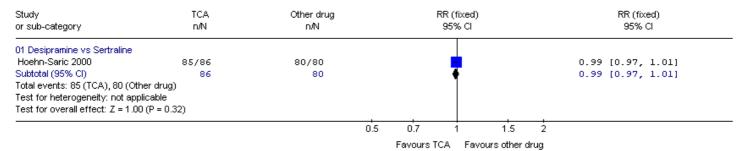
TCAs (OCD)



Review: OCD: TCAs

Comparison: 02 TCAs vs other drugs (adults)

Outcome: 01 Adverse effects



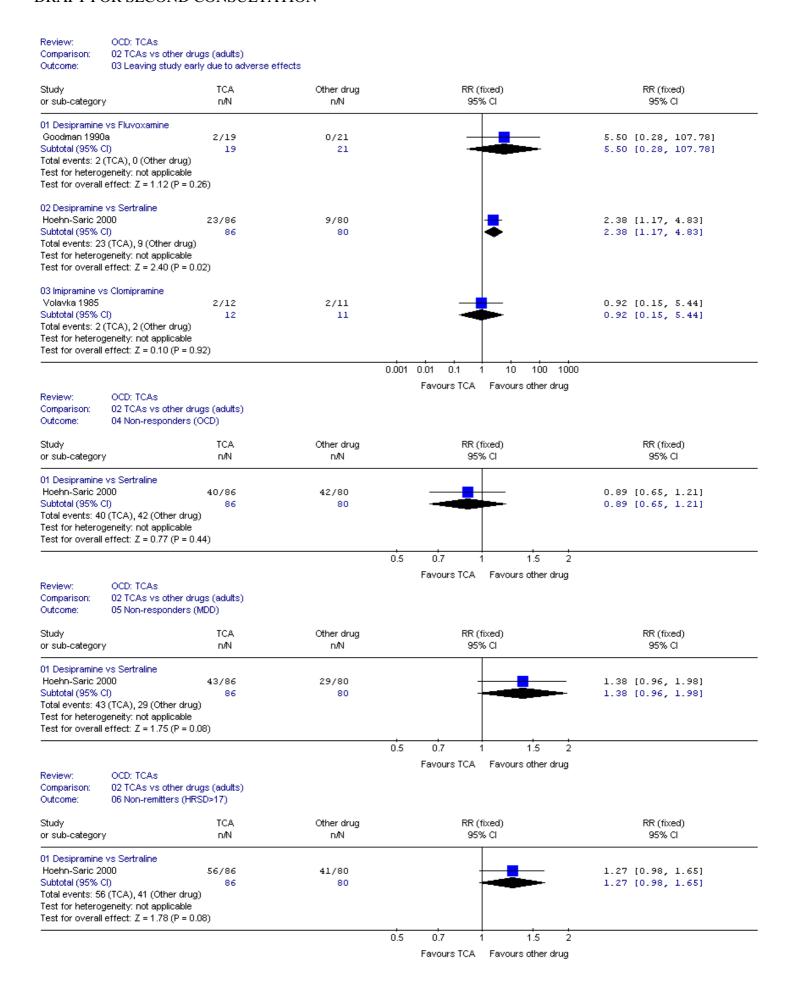
Review: OCD: TCAs

Comparison: 02 TCAs vs other drugs (adults)

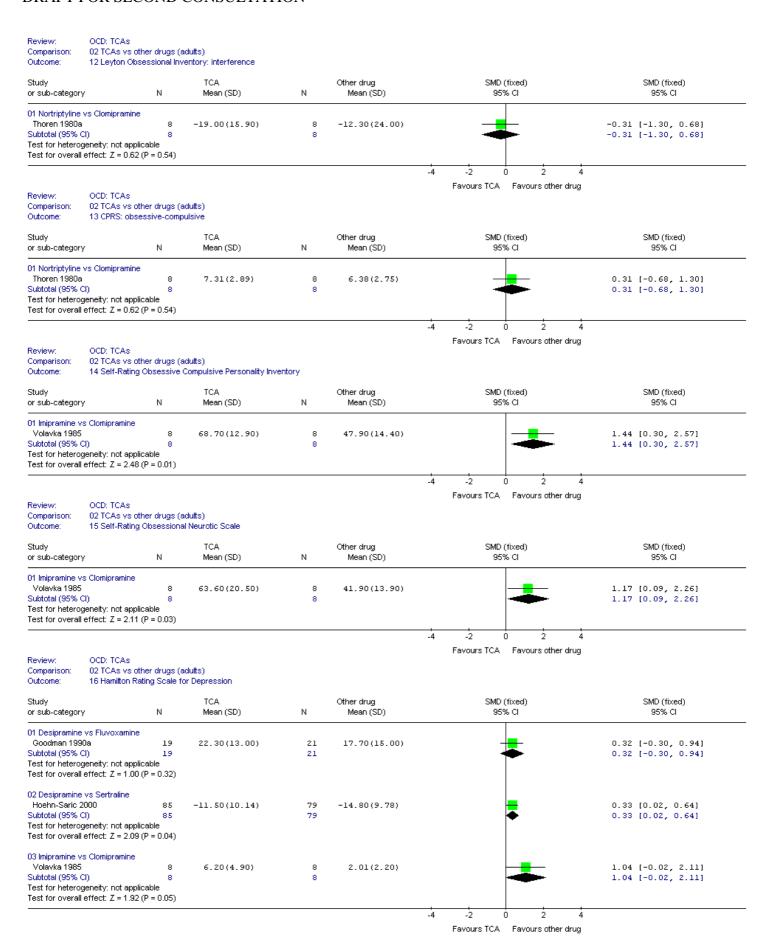
Outcome: 02 Leaving study early

Study or sub-category	TCA n/N	Other drug n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
O1 Amitriptyline vs Clomipramine				
Ananth 1981	2/10	1/10		2.00 [0.21, 18.69]
Subtotal (95% CI)	10	10		2.00 [0.21, 18.69]
Total events: 2 (TCA), 1 (Other drug)				
Test for heterogeneity: not applicable				
Test for overall effect: $Z = 0.61$ (P = 0).54)			
02 Desipramine vs Fluvoxamine				
Goodman 1990a	6/19	2/21	+	3.32 [0.76, 14.49]
Subtotal (95% CI)	19	21		3.32 [0.76, 14.49]
Total events: 6 (TCA), 2 (Other drug)				
Test for heterogeneity: not applicable				
Test for overall effect: Z = 1.59 (P = 0	0.11)			
03 Desipramine vs Sertraline				
Hoehn-Saric 2000	33/86	14/80	-	2.19 [1.27, 3.79]
Subtotal (95% CI)	86	80	◆	2.19 [1.27, 3.79]
Total events: 33 (TCA), 14 (Other dru				
Test for heterogeneity: not applicable				
Test for overall effect: $Z = 2.82$ (P = 0	0.005)			
04 Imipramine vs Clomipramine			L	
Volavka 1985	4/12	3/11		1.22 [0.35, 4.28]
Subtotal (95% CI)	12	11		1.22 [0.35, 4.28]
Total events: 4 (TCA), 3 (Other drug)				
Test for heterogeneity: not applicable				
Test for overall effect: $Z = 0.31$ (P = 0).75)			

Favours TCA Favours other drug

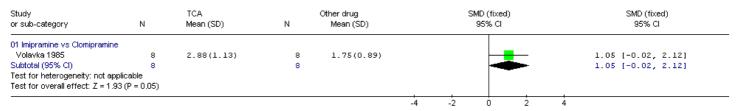


OCD: TCAs Review: Comparison: 02 TCAs vs other drugs (adults) Outcome 07 Y-BOCS SMD (fixed) SMD (fixed) Study TCA Other drug or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Desipramine vs Fluvoxamine Goodman 1990a 25.80(6.00) 21 19.50(9.00) 0.80 [0.15, 1.45] 19 Subtotal (95% CI) 19 21 0.80 [0.15, 1.45] Test for heterogeneity: not applicable Test for overall effect: Z = 2.42 (P = 0.02) Hoehn-Saric 2000 85 -6.00(8.30) 79 -8.40(8.00) 0.29 [-0.02, 0.60] Subtotal (95% CI) 85 79 0.29 [-0.02, 0.60] Test for heterogeneity; not applicable Test for overall effect: Z = 1.86 (P = 0.06) 0.39 [0.11, 0.66] Total (95% CI) 100 Test for heterogeneity: Chi² = 1.92, df = 1 (P = 0.17), l² = 47.8% Test for overall effect: Z = 2.72 (P = 0.006) ń Favours TCA Favours other drug OCD: TCAs Review: Comparison: 02 TCAs vs other drugs (adults) 08 NIMH-OC Outcome SMD (fixed) SMD (fixed) Other drug or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Desipramine vs Sertraline Hoehn-Saric 2000 85 -1.70(2.77) 79 -2.70(2.67) 0.37 [0.06, 0.67] Subtotal (95% CI) 0.37 [0.06, 0.67] Test for heterogeneity: not applicable Test for overall effect: Z = 2.32 (P = 0.02) -0.5 Ö 0.5 Favours TCA Favours other drug Review: OCD: TCAs Comparison: 02 TCAs vs other drugs (adults) Outcome: 09 Leyton Obsessional Inventory: symptom SMD (fixed) SMD (fixed) Study TCA Other drug Ν Mean (SD) N Mean (SD) 95% CI 95% CI or sub-category 01 Nortriptyline vs Clomipramine Thoren 1980a 8 -3.00(3.10) -4.40(9.60) 0.19 [-0.80, 1.17] Subtotal (95% CI) 0.19 [-0.80, 1.17] 8 Test for heterogeneity: not applicable Test for overall effect: Z = 0.37 (P = 0.71) -4 0 Favours TCA Favours other drug Review: OCD: TCAs 02 TCAs vs other drugs (adults) Comparison: 10 Leyton Obsessional Inventory: trait Outcome Study Other drug SMD (fixed) SMD (fixed) Mean (SD) Ν Mean (SD) or sub-category Ν 95% CI 95% CI 01 Nortriptyline vs Clomipramine Thoren 1980a -1.50(1.10) -1.90(2.30) 0.21 [-0.77, 1.19] Subtotal (95% CI) 8 0.21 [-0.77, 1.19] Test for heterogeneity; not applicable Test for overall effect: Z = 0.42 (P = 0.68) ż Ó Favours TCA Favours other drug OCD: TCAs Review: Comparison: 02 TCAs vs other drugs (adults) Outcome 11 Levton Obsessional Inventory: resistance SMD (fixed) Other drug SMD (fixed) or sub-category Ν Mean (SD) N Mean (SD) 95% CI 95% CI 01 Nortriptyline vs Clomipramine Thoren 1980a -4.60(15.90) -13.00(19.60) 0.45 [-0.55, 1.44] Subtotal (95% CI) 0.45 [-0.55, 1.44] Test for heterogeneity: not applicable Test for overall effect: Z = 0.88 (P = 0.38)-2 Ó Favours TCA Favours other drug



Review: OCD: TCAs

Comparison: 02 TCAs vs other drugs (adults)
Outcome: 17 Global Evaluation of Efficacy



Review: OCD: TCAs

Comparison: 03 Desipramine substitution vs Clomipramine continuation (child/adolescent)
Outcome: 01 Leaving the study early (only during the 2-month substitution period)

Study or sub-category	Desipramine n/N	Clomipramine n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
Leonard 1991a	1/10	0/11	-	3.27 [0.15, 72.23]
Total (95% CI) Total events: 1 (Desipramin Test for heterogeneity: not a Test for overall effect: Z = 0	applicable	11		3.27 [0.15, 72.23]
		0.01	0.1 1 10 1	00

Favours Desipramine Favours Clomipramine

Favours Desipramine Favours Clomipramine

Favours TCA Favours other drug

Review: OCD: TCAs

Comparison: 03 Designamine substitution vs Clomipramine continuation (child/adolescent)

Outcome: 02 Relapse (Physician's Relapse Scale)

Leonard 1991a 8/9 2/11 — 4.89 [1.37, 17.49] Total (95% Cl) 9 11 4.89 [1.37, 17.49] Total events: 8 (Desipramine), 2 (Clomipramine) Test for heterogeneity: not applicable Test for overall effect: Z = 2.44 (P = 0.01)	Study or sub-category	Desipramine n/N	Clomipramine n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
Total events: 8 (Desipramine), 2 (Clomipramine) Test for heterogeneity: not applicable	Leonard 1991a	8/9	2/11	-	4.89 [1.37, 17.49]
	Total events: 8 (Desipramino Test for heterogeneity: not a	applicable	11		4.89 [1.37, 17.49]

Review: OCD: TCAs

Comparison: 03 Designamine substitution vs Clomipramine continuation (child/adolescent)
Outcome: 03 Comprehensive Psychopathological Rating Scale: obsessive-compulsive

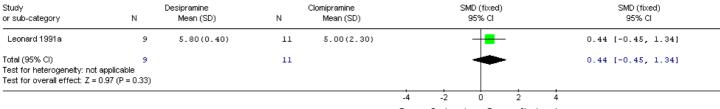
Leonard 1991a 9 6.30 (3.30) 11 5.20 (4.00) — 0.28 [-0.60 Total (95% CI) 9 11 0.28 [-0.60 Test for heterogeneity: not applicable Test for overall effect: Z = 0.63 (P = 0.53)	xed) Cl
Test for heterogeneity: not applicable	, 1.17]
Test for overall effect. Z = 0.05 (P = 0.05)	, 1.17]

Favours Desipramine Favours Clomipramine

Review: OCD: TCAs

Comparison: 03 Designamine substitution vs Clomipramine continuation (child/adolescent)

Outcome: 04 NIMH-OC



Favours Desipramine Favours Clomipramine

SNRIs (OCD)

Review: OCD: SNRIs Comparison: 01 SNRIs vs other drugs 01 Adverse effects Outcome: SNRI RR (fixed) RR (fixed) Study Other drug or sub-category n/Ν n/Ν 95% CI 95% CI 01 Venlafaxine vs Clomipramine 0.67 [0.49, 0.92] Albert 2002 16/26 43/47 Subtotal (95% CI) 0.67 [0.49, 0.92] 26 47 Total events: 16 (SNRI), 43 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 2.46 (P = 0.01) Favours SNRI Favours other drug Review: OCD: SNRIs Comparison: 01 SNRIs vs other drugs 02 Leaving study early Outcome: Other drug RR (fixed) **SNRI** RR (fixed) Study 95% CI or sub-category n/Ν n/N 95% CI 01 Venlafaxine vs Clomipramine Albert 2002 0.26 [0.03, 1.99] 1/26 7/47 Subtotal (95% CI) 0.26 [0.03, 1.99] 26 47 Total events: 1 (SNRI), 7 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 1.30 (P = 0.19)0.01 0.1 10 100 Favours SNRI Favours other drug OCD: SNRIs Review: 01 SNRIs vs other drugs Comparison: Outcome: 03 Leaving study early due to adverse effects Study SNRI Other drug RR (fixed) RR (fixed) 95% CI 95% CI or sub-category nΝ 01 Venlafaxine vs Paroxetine 0.33 [0.07, 1.60] Denvs 2003a 2/75 6/75 Subtotal (95% CI) 75 75 0.33 [0.07, 1.60] Total events: 2 (SNRI), 6 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 1.37 (P = 0.17) 02 Venlafaxine vs Clomipramine Albert 2002 0/26 5/47 0.16 [0.01, 2.81] Subtotal (95% CI) 0.16 [0.01, 2.81] Total events: 0 (SNRI), 5 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 1.25 (P = 0.21) 100 1000 0.001 0.01 0.1 Favours SNRI Favours other drug Review: OCD: SNRIs 01 SNRIs vs other drugs Comparison: 04 Non-responders Outcome: SNRI RR (fixed) RR (fixed) Study Other drug or sub-category n/N n/N 95% CI 95% CI 01 Venlafaxine vs Paroxetine Denys 2003a 47/75 42/75 1.12 [0.86, 1.46] Subtotal (95% CI) 1.12 [0.86, 1.46] 75 75 Total events: 47 (SNRI), 42 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.83 (P = 0.41) 02 Venlafaxine vs Clomipramine 1.14 [0.78, 1.65] Albert 2002 17/26 27/47 Subtotal (95% CI) 26 47 1.14 [0.78, 1.65] Total events: 17 (SNRI), 27 (Other drug) Test for heterogeneity: not applicable

Favours SNRI Favours other drug

Test for overall effect: Z = 0.68 (P = 0.50)

Review: OCD: SNRIs

01 SNRIs vs other drugs 05 Y-BOCS Comparison:

Outcome:

Study or sub-category	N	SNRI Mean (SD)	N	Other drug Mean (SD)		SMD (fi 95%	*	SMD (fixed) 95% Cl
01 Venlafaxine vs Paroxetin	ne							
Denys 2003a	73	-7.20(7.50)	72	-7.80(5.40)				0.09 [-0.23, 0.42]
Subtotal (95% CI)	73		72					0.09 [-0.23, 0.42]
Test for heterogeneity: not a Test for overall effect: Z = 0							_	
02 Venlafaxine vs Clomiprar	mine							
Albert 2002	26	18.36(7.11)	47	17.30(6.15)				0.16 [-0.32, 0.64]
Subtotal (95% CI)	26		47					0.16 [-0.32, 0.64]
Test for heterogeneity: not a	applicable							
Test for overall effect: $Z = 0$	0.66 (P = 0.51)							
					-1	-0.5 0	0.5	1
					F	avours SNRI	Favours other drug	9

Review: OCD: SNRIs

Comparison:

01 SNRIs vs other drugs 06 Hamilton Rating Scale for Depression Outcome:

(fixed) % Cl
19, 0.46]
19, 0.46]
_

Favours SNRI Favours other drug

Review: OCD: SNRIs

Comparison: 01 SNRIs vs other drugs Outcome: 07 Hamilton Anxiety Scale

Study or sub-category	N	SNRI Mean (SD)	N	Other drug Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Venlafaxine vs Paroxet	ine					
Denys 2003a	73	-4.90(5.00)	72	-4.70(7.00)		-0.03 [-0.36, 0.29]
Subtotal (95% CI)	73		72			-0.03 [-0.36, 0.29]
Test for heterogeneity: not	: applicable					
Test for overall effect: Z =	0.20 (P = 0.84)					
					-0.5 -0.25 0 0.25 0.5	
					Favours SNRI Favours other drug	

MAOIs (OCD)

OCD: MAOIs Review: 01 MAOIs vs placebo Comparison: Outcome: 01 Leaving study early

Study or sub-category	MAOI n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Phenelzine vs Placebo				
Jenike 1997	3/20	3/21		— 1.05 [0.24, 4.61]
Subtotal (95% CI)	20	21		- 1.05 [0.24, 4.61]
Total events: 3 (MAOI), 3 (Place	cebo)			•
Test for heterogeneity: not ap				
Test for overall effect: $Z = 0.0$	06 (P = 0.95)			
		0.2	0.5 1 2	5
			Favours MAOI Favours placeb	00

Review: OCD: MAOIs 01 MAOIs vs placebo 02 Y-BOCS Comparison: Outcome:

Study or sub-category	N	MAOI Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl					SMD (fixed) 95% Cl
01 Phenelzine vs Placebo Jenike 1997	1.7	16 0048 001	10	10 7046 101						0.05 / 1.00 0.011
Subtotal (95% CI)	17 17	16.30(7.00)	18 18	18.70(6.10)		-	•			-0.36 [-1.03, 0.31] -0.36 [-1.03, 0.31]
Test for heterogeneity: not Test for overall effect: Z =										
					-4	-2	0	2	4	
					Favours MAOI Favours placebo					

OCD: MAOIs Review: Comparison: 01 MAOIs vs placebo Outcome 03 NIMH-OC SMD (fixed) SMD (fixed) Study MAOI Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Phenelzine vs Placebo Jenike 1997 17 6.80(1.40) 18 7.20(1.30) -0.29 [-0.96, 0.38] Subtotal (95% CI) -0.29 [-0.96, 0.38] Test for heterogeneity: not applicable Test for overall effect: Z = 0.85 (P = 0.39) -0.5 ò 0.5 Favours MAOI Favours placebo OCD: MAOIs Review: 01 MAOIs vs placebo Comparison: Outcome 04 OCD Scale (CPRS) SMD (fixed) Study SMD (fixed) MAOL Placebo Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI or sub-category 01 Phenelzine vs Placebo Jenike 1997 17 6.50(3.80) 18 5.80(3.30) 0.19 [-0.47, 0.86] Subtotal (95% CI) 0.19 [-0.47, 0.86] 18 17 Test for heterogeneity: not applicable Test for overall effect: Z = 0.57 (P = 0.57) -0.5 0.5 Ó Favours MAOI Favours placebo Review: OCD: MAOIs 01 MAOIs vs placebo Comparison: 05 Clinical Global Impression Outcome Study MAOI Placebo SMD (fixed) SMD (fixed) Mean (SD) Ν or sub-category Ν Mean (SD) 95% CI 95% CI 01 Phenelzine vs Placebo Jenike 1997 3.20(1.00) 18 3.20(1.00) 0.00 [-0.66, 0.66] Subtotal (95% CI) 17 18 0.00 [-0.66, 0.66] Test for heterogeneity; not applicable Test for overall effect: Z = 0.00 (P = 1.00) -0.5 ń 0.5 Favours MAOI Favours placebo Review: OCD: MAOIs 02 MAOIs vs other drugs Comparison: Outcome: 01 Leaving study early Study MAOI Other drug RR (fixed) RR (fixed) 95% CI or sub-category n/N n/N 95% CI 01 Phenelzine vs Clomipramine Valleio 1992 2/14 2/16 1.14 [0.18, 7.08] Subtotal (95% CI) 16 1.14 [0.18, 7.08] Total events: 2 (MAOI), 2 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.14 (P = 0.89) 02 Phenelzine vs Fluoxetine Jenike 1997 3/20 4/23 0.86 [0.22, 3.40] Subtotal (95% CI) 23 0.86 [0.22, 3.40] 20 Total events: 3 (MAOI), 4 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.21 (P = 0.83) 0.1 0.2 0.5 5 10 Favours MAOL Favours other drug Review: OCD: MAOIs Comparison: 02 MAOIs vs other drugs 02 Leaving study early due to adverse effects Outcome: Study MAOI Other drug RR (fixed) RR (fixed) or sub-category nNnN95% CI 95% CI 01 Phenelzine vs Clomipramine Vallejo 1992 0/14 1/16 0.38 [0.02, 8.59] Subtotal (95% CI) 14 16 0.38 [0.02, 8.59] Total events: 0 (MAOI), 1 (Other drug) Test for heterogeneity: not applicable Test for overall effect: Z = 0.61 (P = 0.54) 0.01

Favours MAOI Favours other drug

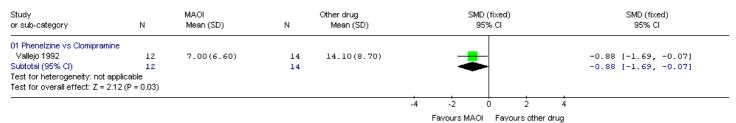
OCD: MAOIs Review: Comparison: 02 MAOIs vs other drugs Outcome 03 Y-BOCS SMD (fixed) SMD (fixed) Study MAOI Other drug or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Phenelzine vs Fluoxetine Jenike 1997 17 16.30(7.00) 19 16.20(6.30) 0.01 [-0.64, 0.67] Subtotal (95% CI) 19 0.01 [-0.64, 0.67] Test for heterogeneity: not applicable Test for overall effect: Z = 0.04 (P = 0.96) -0.5 ò 0.5 Favours MAOI Favours other drug OCD: MAOIs Review: 02 MAOIs vs other drugs Comparison: Outcome 04 NIMH-OC SMD (fixed) SMD (fixed) Study MAOI Other drug or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Phenelzine vs Fluoxetine -0.16 [-0.81, 0.50] -0.16 [-0.81, 0.50] Jenike 1997 17 6.80(1.40) 19 7.00(1.10) Subtotal (95% CI) 19 17 Test for heterogeneity: not applicable Test for overall effect: Z = 0.47 (P = 0.64) -0.5 0.5 Ó Favours MAOI Favours other drug Review: OCD: MAOIs 02 MAOIs vs other drugs Comparison: Outcome: 05 Maudsley Obsessive-Compulsive Inventory Study MAOL Other drug SMD (fixed) SMD (fixed) Ν Mean (SD) Ν Mean (SD) or sub-category 95% CI 95% CI 01 Phenelzine vs Clomipramine Vallejo 1992 12 10.10(6.90) 12.10(4.70) -0.33 [-1.11, 0.44] Subtotal (95% CI) 12 14 -0.33 [-1.11, 0.44] Test for heterogeneity; not applicable Test for overall effect: Z = 0.84 (P = 0.40) -4 -2 ń Favours MAOI Favours other drug OCD: MAOIs Review: Comparison: 02 MAOIs vs other drugs Outcome: 06 OCD Scale (CPRS) SMD (fixed) SMD (fixed) Study MAOI Other drug or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Phenelzine vs Fluoxetine Jenike 1997 -0.06 [-0.72, 0.59] 6.50(3.80) 19 6.70(2.60) Subtotal (95% CI) -0.06 [-0.72, 0.59] Test for heterogeneity: not applicable Test for overall effect: Z = 0.18 (P = 0.86) -0.5 0.5 Favours MAOI Favours other drug Review: OCD: MAOIs Comparison: 02 MAOIs vs other drugs Outcome 07 Hamilton Rating Scale for Depression SMD (fixed) SMD (fixed) Study MAOI Other drug or sub-category Mean (SD) Ν Mean (SD) 95% CI 95% CI Ν 01 Phenelzine vs Clomipramine Valleio 1992 7.00(5.70) 14 11.50(6.90) -0.68 [-1.48, 0.11] Subtotal (95% CI) -0.68 [-1.48, 0.11] 12 14 Test for heterogeneity: not applicable Test for overall effect: Z = 1.68 (P = 0.09)

-2 0 2 4
Favours MAOI Favours other drug

Review: OCD: MAOIs

Comparison: 02 MAOIs vs other drugs

Outcome: 08 Hamilton Rating Scale for Anxiety



Review: OCD: MAOIs

Comparison: 02 MAOIs vs other drugs Outcome: 09 Clinical Global Impression

Study or sub-category	N	MAOI Mean (SD)	N	Other drug Mean (SD)			MD (fixed 95% Cl)		SMD (fixed) 95% Cl
01 Phenelzine vs Fluoxetine	•									
Jenike 1997	17	3.20(1.00)	19	3.10(1.00)			_			0.10 [-0.56, 0.75]
Subtotal (95% CI)	17		19							0.10 [-0.56, 0.75]
Test for heterogeneity: not	applicable						1-			
Test for overall effect: $Z = 0$	0.29 (P = 0.77)									
					-1	-0.5	-	0.5	1	
					F	avours MA	OI Fav	ours other	drug	

Anxiolytics (OCD)

Review: OCD: anxiolytics

Comparison: 01 Anxiolytics vs placebo Outcome: 01 Leaving study early

Study or sub-category	Anxiolytic n/N	Placebo n/N	RR (fixed) 95% Cl		RR (fixed) 95% Cl
01 Clonazepam					
Hollander 2003c	5/17	4/10			0.74 [0.26, 2.12]
Subtotal (95% CI)	17	10		- -	0.74 [0.26, 2.12]
Total events: 5 (Anxiolytic),	4 (Placebo)				
Test for heterogeneity: not a	applicable				
Test for overall effect: Z = 0	.57 (P = 0.57)				
			0.2 0.5 1	2 5	

Favours anxiolytic

Favours placebo

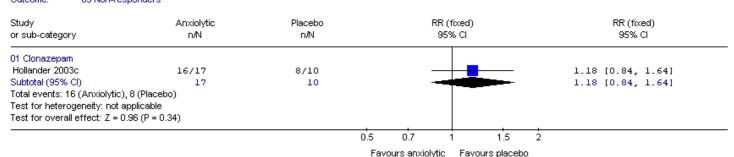
Review: OCD: anxiolytics

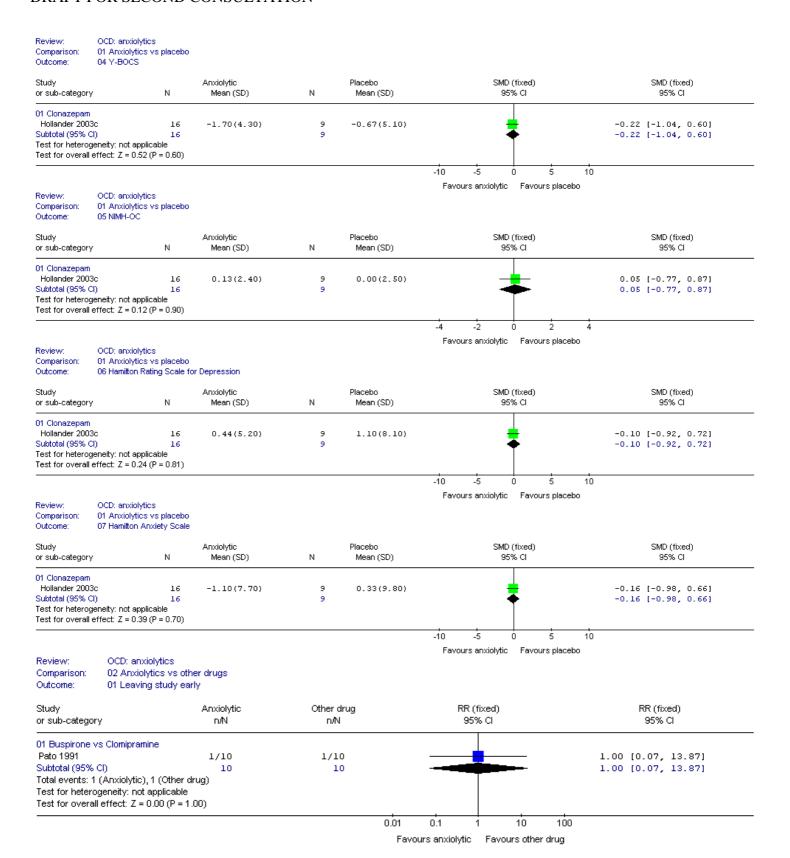
Comparison: 01 Anxiolytics vs placebo

Outcome: 02 Leaving study early due to adverse effects

Study or sub-category	Anxiolytic n/N	Placebo n/N	RR (fixed) RR (fixed) 95% Cl 95% Cl
01 Clonazepam			
Hollander 2003c	3/17	2/10	0.88 [0.18, 4.41]
Subtotal (95% CI)	17	10	0.88 [0.18, 4.41]
Total events: 3 (Anxiolytic), 2	2 (Placebo)		
Test for heterogeneity: not a	pplicable		
Test for overall effect: $Z = 0$.	.15 (P = 0.88)		
			0.1 0.2 0.5 1 2 5 10
			Favours anxiolytic Favours placebo

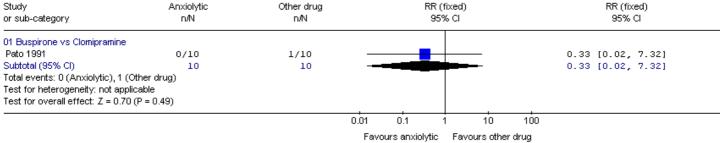
Review: OCD: anxiolytics Comparison: 01 Anxiolytics vs placebo Outcome: 03 Non-responders





OCD: anxiolytics Review: 02 Anxiolytics vs other drugs Comparison: Outcome:

02 Leaving study early due to adverse effects



OCD: anxiolytics Review:

Comparison: 02 Anxiolytics vs other drugs

Outcome: 03 Y-BOCS

Study or sub-category	N	Anxiolytic Mean (SD)	N	Other drug Mean (SD)		S	SMD (fixed) 95% CI			SMD (fixed) 95% Cl
01 Buspirone vs Clomipran	nine									
Pato 1991	9	16.78(6.46)	9	17.67(4.97)			-			-0.15 [-1.07, 0.78]
Subtotal (95% CI)	9		9				•			-0.15 [-1.07, 0.78]
Test for heterogeneity: not	: applicable						I			·
Test for overall effect: Z =	0.31 (P = 0.76)									
					-10	-5	-	5	10	

Favours anxiolytic Favours other drug

Favours anxiolytic Favours other drug

Review: OCD: anxiolytics

02 Anxiolytics vs other drugs 04 NIMH-OC Comparison:

Outcome:

Study or sub-category	N	Anxiolytic Mean (SD)	N	Other drug Mean (SD)			MD (fixed) 95% Cl			SMD (fixed) 95% Cl
01 Buspirone vs Clomipran	nine									
Pato 1991	9	5.11(1.27)	9	5.22(1.20)		_	-		-	0.08 [-1.01, 0.84]
Subtotal (95% CI)	9		9			-4	•		_	0.08 [-1.01, 0.84]
Test for heterogeneity: not	: applicable						Τ			
Test for overall effect: Z =	0.18 (P = 0.86)									
					-4				4	

OCD: anxiolytics Review:

Comparison: 02 Anxiolytics vs other drugs

Outcome: 05 Hamilton Rating Scale for Depression

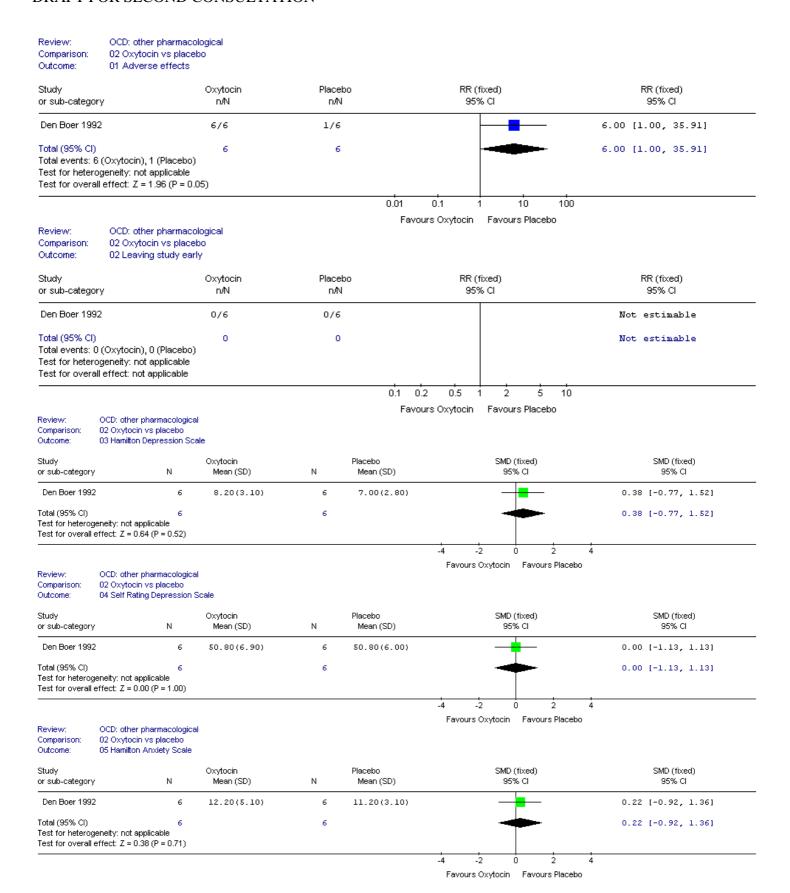
Study or sub-category	N	Anxiolytic Mean (SD)	N	Other drug Mean (SD)		5	SMD (fixed 95% Cl))		SMD (fixed) 95% Cl
01 Buspirone vs Clomipramine)									
Pato 1991	9	9.89(6.75)	9	9.33(4.58)			-			0.09 [-0.83, 1.02]
Subtotal (95% CI)	9		9				•			0.09 [-0.83, 1.02]
Test for heterogeneity: not ap	plicable						ſ			•
Test for overall effect: $Z = 0.2$	20 (P = 0.84)									
					-10	-5	-	5	10	
					Favo	ours anxiol	ytic Fav	ours othe	r drug	

Other Pharmacological (OCD)

Review: OCD: other pharmacological 01 Inositol vs placebo Comparison:

Outcome 01 Y-BOCS

Study or sub-category	N	Inositol Mean (SD)	N	Placebo Mean (SD)			1D (fixed) 95% CI			SMD (fixed) 95% Cl
Fux 1996	7	17.57(9.60)	6	20.00(5.20)		_	-			-0.29 [-1.38, 0.81]
Total (95% CI) Test for heterogeneity: not			6				-			-0.29 [-1.38, 0.81]
Test for overall effect: Z =	0.51 (P = 0.61)									
					-4	-2	Ó	2	4	
					Far	vours Inosit	ol Favo	ours Place	ebo	

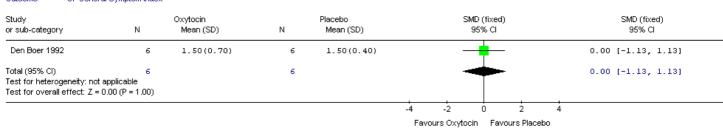


Review: OCD: other pharmacological Comparison: 02 Oxytocin vs placebo Outcome: 06 State Anxiety Inventory

Study or sub-category	N	Oxytocin Mean (SD)	N	Placebo Mean (SD)			MD (fixed) 95% Cl	ı		SMD (fixed) 95% Cl
Den Boer 1992	6	48.70(13.40)	6	61.20(7.90)		-	_			-1.05 [-2.29, 0.19]
Total (95% Cl) Test for heterogeneity: not Test for overall effect: Z =			6							-1.05 [-2.29, 0.19]
					-4	-2	ò	2	4	

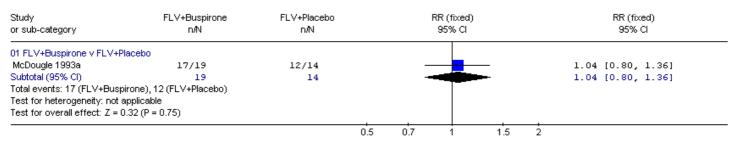
Favours Oxytocin Favours Placebo

Review: OCD: other pharmacological Comparison: 02 Oxytocin vs placebo Outcome: 07 General Symptom Index



Augmentation (OCD)

Review: OCD: augmentation strategies
Comparison: 01 Buspirone vs placebo
Outcome: 01 Non-responders



Favours augmentation Favours non-aug

Favours augmentation Favours non-aug

Review: OCD: augmentation strategies Comparison: 01 Buspirone vs placebo

Outcome: 02 Y-BOCS

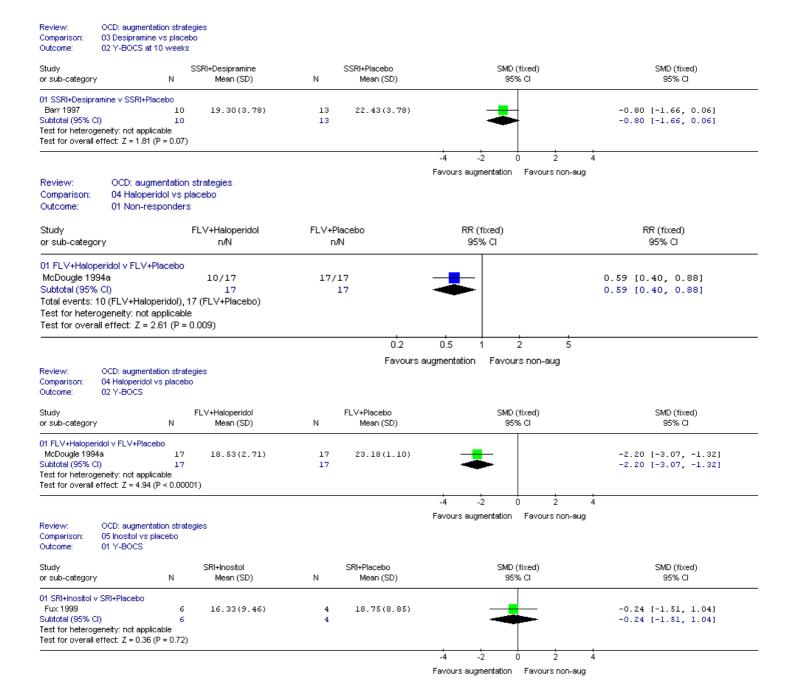
Study or sub-category	N	FLV+Buspirone Mean (SD)	N	FLV+Placebo Mean (SD)			MD (fixed) 95% Cl			SMD (fixed) 95% Cl
01 FLV+Buspirone v FLV+ McDougle 1993a Subtotal (95% CI) Test for heterogeneity: not Test for overall effect: Z =	19 19 t applicable	24.90(6.60)	14 14	21.00(9.10)						0.49 [-0.21, 1.19] 0.49 [-0.21, 1.19]
					-4	-2	0	2	4	

Review: OCD: augmentation strategies
Comparison: 01 Buspirone vs placebo
Outcome: 03 Hamilton Depression Rating Scale

Study or sub-category	N	FLV+Buspirone Mean (SD)	N	FLV+Placebo Mean (SD)			MD (fixed) 95% Cl			SMD (fixed) 95% Cl
01 FLV+Buspirone v FLV+PI McDougle 1993a Subtotal (95% CI) Test for heterogeneity: not a Test for overall effect: Z = 1	19 19 applicable	17.90(10.70)	14 14	11.70(7.40)			*			0.64 [-0.07, 1.35] 0.64 [-0.07, 1.35]
					-4	-2 augmentatio	0 D Favou	2 rs non-au	4	

OCD: augmentation strategies Review: Comparison: 01 Buspirone vs placebo Outcome 04 Hamilton Anxiety Scale FLV+Placebo SMD (fixed) SMD (fixed) Study FLV+Buspirone or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 FLV+Buspirone v FLV+Placebo McDougle 1993a 11.60(6.80) 8.60(4.90) 0.48 [-0.22, 1.18] 19 14 Subtotal (95% CI) 19 0.48 [-0.22, 1.18] Test for heterogeneity: not applicable Test for overall effect: Z = 1.35 (P = 0.18) ò Favours augmentation Favours non-aug OCD: augmentation strategies Review: Comparison: 02 Citalopram+Clomipramine vs Citalopram Outcome: 01 Non-responders (Y-BOCS 35%) Study Citalopram+CMI Citalopram RR (fixed) RR (fixed) 95% CI 95% CI or sub-category n/N n/N Pallanti 1999 0/9 6/7 0.06 [0.00, 0.94] Total (95% CI) 0.06 [0.00, 0.94] Total events: 0 (Citalopram+CMI), 6 (Citalopram) Test for heterogeneity: not applicable Test for overall effect: Z = 2.01 (P = 0.04) 0.001 0.01 0.1 10 100 1000 Favours augmentation Favours non-aug OCD: augmentation strategies Comparison: 02 Citalopram+Clomipramine vs Citalopram 02 Y-BOCS Outcome Citalopram+CMI Study Citalopram SMD (fixed) SMD (fixed) or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI Pallanti 1999 9 25.60(4.80) 15.90(3.80) -2.15 [-3.46, -0.84] Total (95% CI) -2.15 [-3.46, -0.84] Test for heterogeneity: not applicable Test for overall effect: Z = 3.22 (P = 0.001) Ö Favours augmentation Favours non-aug OCD: augmentation strategies Review: Comparison: 02 Citalopram+Clomipramine vs Citalopram Outcome 03 Hamilton Depression Rating Scale SMD (fixed) SMD (fixed) Study Citalopram+CMI Citalopram N Mean (SD) 95% CI or sub-category Ν Mean (SD) Pallanti 1999 9 5.60(2.60) 6.40(1.80) -0.33 [-1.33, 0.67] Total (95% CI) -0.33 [-1.33, 0.67] Test for heterogeneity: not applicable Test for overall effect: Z = 0.65 (P = 0.52) 0 Favours augmentation Favours non-aug Review: OCD: augmentation strategies 03 Desipramine vs placebo Comparison: 01 Y-BOCS at 6 weeks Outcome Study SSRI+Desipramine SSRI+Placebo SMD (fixed) SMD (fixed) or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 SSRI+Desipramine v SSRI+Placebo Barr 1997 18.52(2.94) 13 24.52(2.94) -1.97 [-3.00, -0.93] Subtotal (95% CI) 10 13 -1.97 [-3.00, -0.93] Test for heterogeneity: not applicable Test for overall effect: Z = 3.73 (P = 0.0002)

-4 -2 0 2
Favours augmentation Favours non-aug



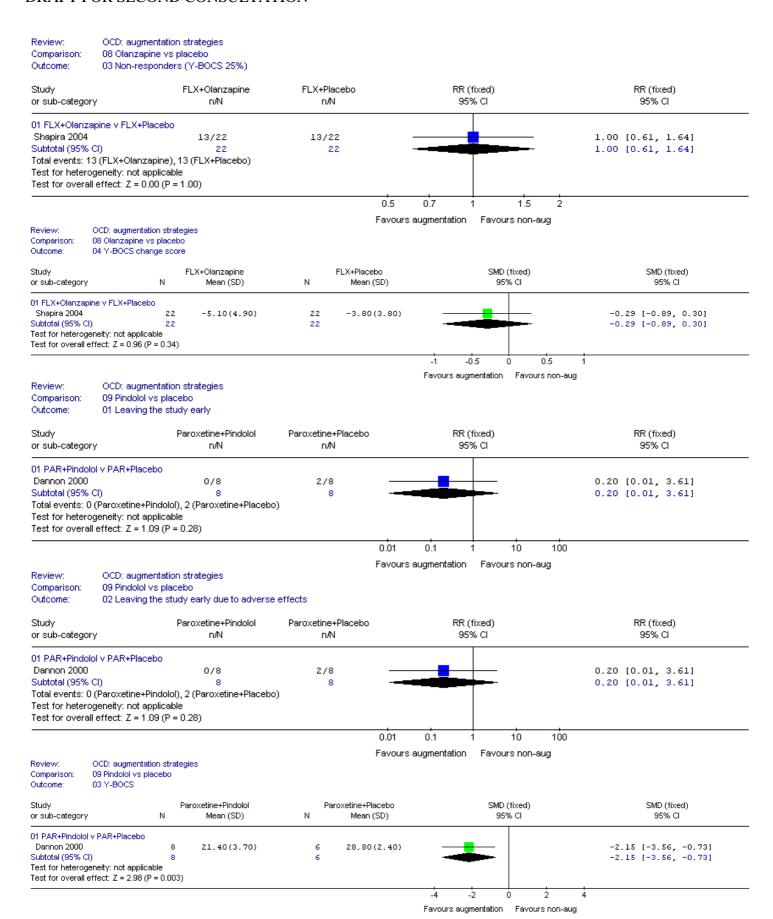
Review: OCD: augmentation strategies 06 Lithium vs placebo Comparison: 01 Non-responders Outcome: Study FLV+Lithium FLV+Placebo RR (fixed) RR (fixed) n/N 95% CI 95% CI or sub-category n/N 01 FLV+Lithium v FLV+Placebo at 2 weeks McDougle 1991 (1) 9/11 9/9 0.82 [0.62, 1.08] Subtotal (95% CI) 9 0.82 [0.62, 1.08] Total events: 9 (FLV+Lithium), 9 (FLV+Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.41 (P = 0.16) 02 FLV+Lithium v FLV+Placebo at 4 weeks McDougle 1991 (2) 5/5 Not estimable Subtotal (95% CI) Not estimable Total events: 0 (FLV+Lithium), 0 (FLV+Placebo) Test for heterogeneity: not applicable Test for overall effect: not applicable Total (95% CI) 16 14 0.82 [0.62, 1.08] Total events: 14 (FLV+Lithium), 14 (FLV+Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.41 (P = 0.16) 0.5 0.7 1.5 Favours augmentation Favours non-aug Review: OCD: augmentation strategies Comparison: 06 Lithium vs placebo 02 Y-BOCS change score Outcome Study FLV+Lithium FLV+Placebo SMD (fixed) SMD (fixed) Ν Mean (SD) Ν Mean (SD) 95% CI or sub-category 95% CI 01 FLV+Lithium v FLV+Placebo at 2 weeks McDougle 1991 (1) 11 -4.00(5.23) 1.22(5.24) -0.96 [-1.90, -0.01] Subtotal (95% CI) -0.96 [-1.90, -0.01] 11 Test for heterogeneity: not applicable Test for overall effect: Z = 1.99 (P = 0.05) 02 FLV+Lithium v FLV+Placebo at 4 weeks McDougle 1991 (2) 2.60(3.58) -1.40(2.07) 1.24 [-0.19, 2.66] 5 1.24 [-0.19, 2.66] Subtotal (95% CI) Test for heterogeneity: not applicable Test for overall effect: Z = 1.70 (P = 0.09) -0.29 [-1.07, 0.50] Test for heterogeneity: Chi² = 6.34, df = 1 (P = 0.01), l² = 84.2% Test for overall effect: Z = 0.72 (P = 0.47) Favours augmentation Favours non-aug OCD: augmentation strategies Review: Comparison: 06 Lithium vs placebo Outcome 03 Hamilton Depression Scale change score FLV+Placebo FLV+Lithium SMD (fixed) SMD (fixed) Study or sub-category N 95% CI 95% CI Mean (SD) Mean (SD) 01 FLV+Lithium v FLV+Placebo at 2 weeks McDougle 1991 (1) 1.1 -7.0078.811 0.00(9.06) -0.75 [-1.67, 0.17] -0.75 [-1.67, 0.17] Subtotal (95% CI) 11 Test for heterogeneity: not applicable Test for overall effect: Z = 1.60 (P = 0.11) 02 FLV+Lithium v FLV+Placebo at 4 weeks McDougle 1991 (2) 4.60(6.15) -3.40(4.28) 1.36 [-0.09, 2.82] Subtotal (95% CI) 1.36 [-0.09, 2.82] Test for heterogeneity: not applicable Test for overall effect: Z = 1.83 (P = 0.07) Total (95% CI) -0.15 [-0.93, 0.63] Test for heterogeneity: $Chi^2 = 5.79$, df = 1 (P = 0.02), $I^2 = 82.7\%$ Test for overall effect: Z = 0.38 (P = 0.70)

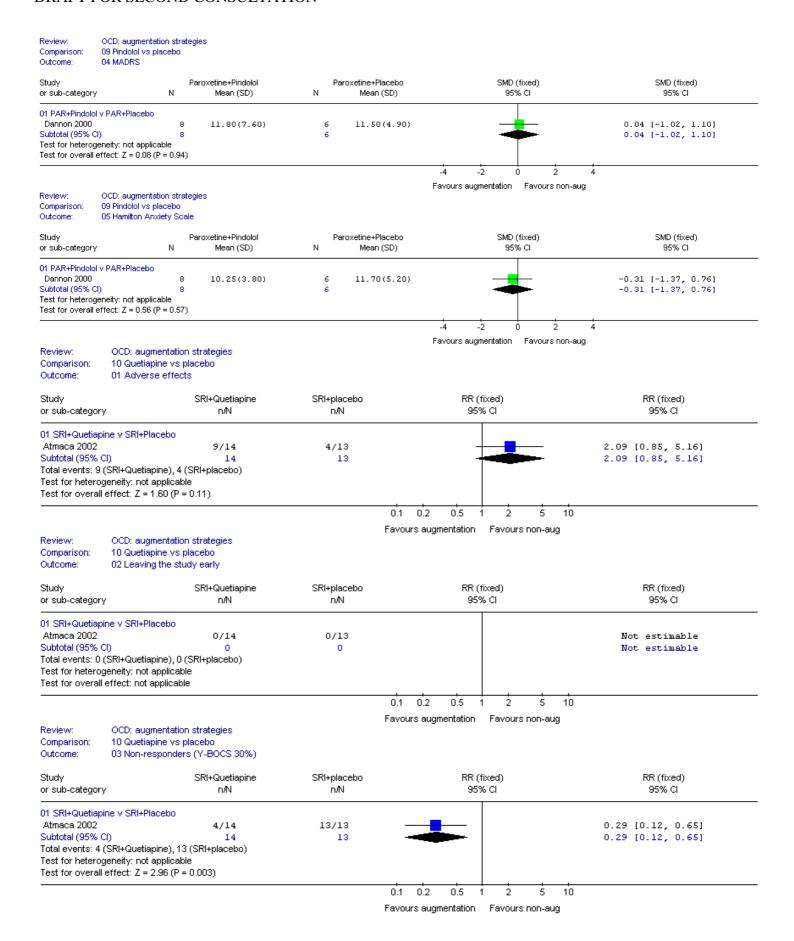
Favours augmentation Favours non-aug

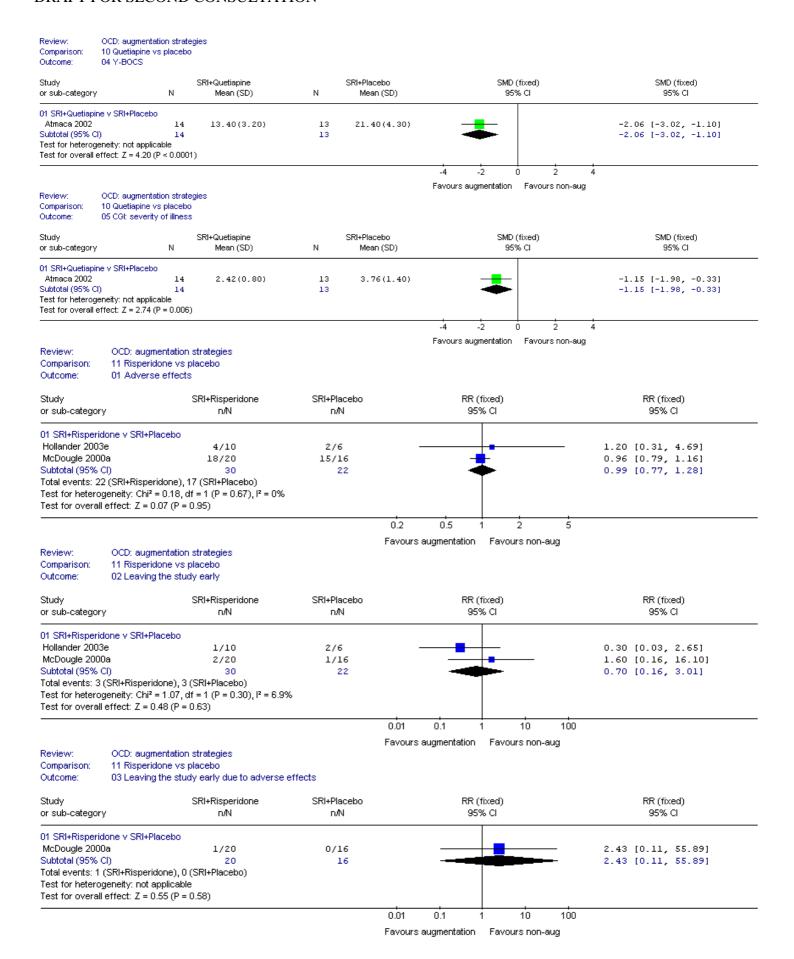
OCD: augmentation strategies Review: Comparison: 06 Lithium vs placebo Outcome 04 Hamilton Anxiety Scale change score FLV+Placebo SMD (fixed) SMD (fixed) Study or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 FLV+Lithium v FLV+Placebo at 2 weeks McDougle 1991 (1) -0.73(4.67) -1.11(3.26) 0.09 [-0.79, 0.97] 11 Subtotal (95% CI) 11 0.09 [-0.79, 0.97] Test for heterogeneity: not applicable Test for overall effect: Z = 0.20 (P = 0.84) 02 FLV+Lithium v FLV+Placebo at 4 weeks McDougle 1991 (2) Subtotal (95% CI) 1.40(3.51) -3.20(3.27) 1.22 [-0.19, 2.64] 5 1.22 [-0.19, 2.64] Test for heterogeneity; not applicable Test for overall effect: Z = 1.69 (P = 0.09) 0.41 [-0.34, 1.15] Total (95% CI) 14 Test for heterogeneity: Chi² = 1.78, df = 1 (P = 0.18), l² = 43.8% Test for overall effect: Z = 1.06 (P = 0.29) ń Favours augmentation Favours non-aug Review: OCD: augmentation strategies Comparison: 07 Nortriptyline vs placebo Outcome 01 Y-BOCS CMI+Placebo SMD (fixed) SMD (fixed) CMI+Nortriptyline or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 CMI+Nortriptyline v CMI+Placebo Noorbala 1998 15 8.33(5.56) 15 19.17(5.56) -1.90 [-2.78, -1.02] Subtotal (95% CI) 15 -1.90 [-2.78, -1.02] Test for heterogeneity; not applicable Test for overall effect: Z = 4.22 (P < 0.0001)Ö Favours augmentation Favours non-aug OCD: augmentation strategies Review: Comparison: 08 Olanzapine vs placebo Outcome: 01 Leaving the study early FLX+Placebo Study FLX+Olanzapine RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 FLX+Olanzapine v FLX+Placebo Shapira 2004 5/22 2/22 2.50 [0.54, 11.54] Subtotal (95% CI) 2.50 [0.54, 11.54] Total events: 5 (FLX+Olanzapine), 2 (FLX+Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.17 (P = 0.24) 0.01 0.1 10 100 Favours augmentation Favours non-aug OCD: augmentation strategies Review: 08 Olanzapine vs placebo Comparison: 02 Leaving the study early due to adverse effects Outcome: Study FLX+Olanzapine FLX+Placebo RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 FLX+Olanzapine v FLX+Placebo 2/22 1.00 [0.15, 6.48] Shapira 2004 2/22 Subtotal (95% CI) 22 22 1.00 [0.15, 6.48] Total events: 2 (FLX+Olanzapine), 2 (FLX+Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.00 (P = 1.00) 0.2 0.5 5 10

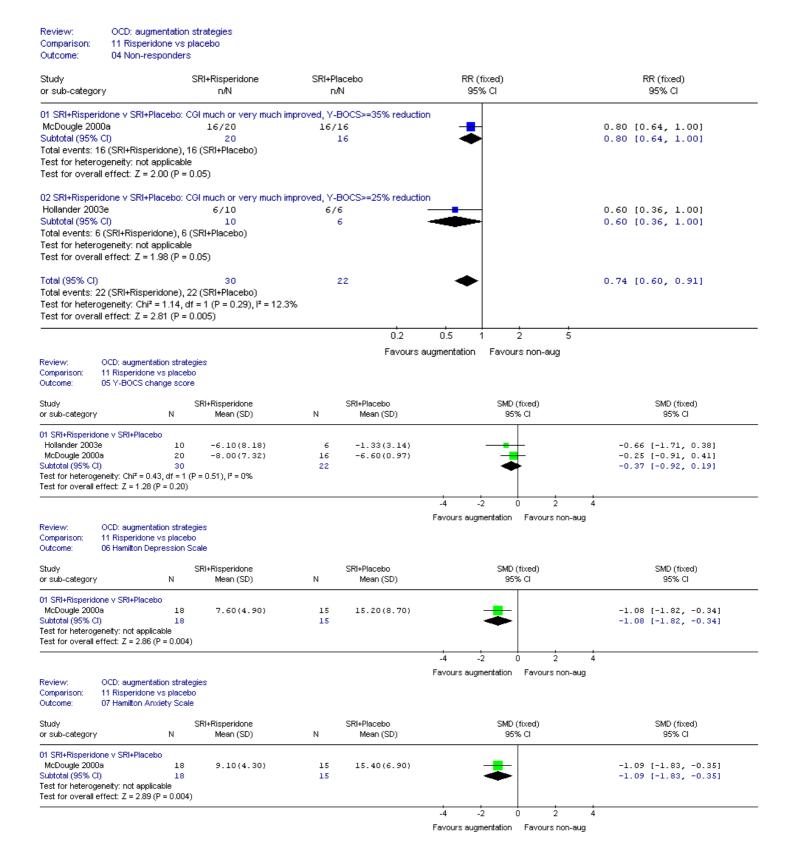
Favours augmentation

Favours non-aug









Review: OCD: augmentation strategies Comparison: 12 All antipsychotics v placebo
Outcome: 01 Adverse effects

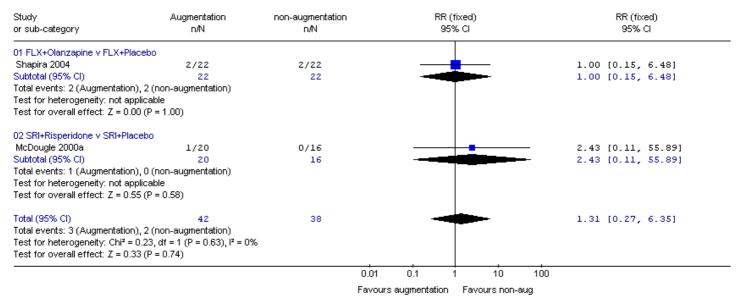
Study or sub-category	Augmentation n/N	non-augmentation n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 SRI+Quetiapine v SRI+Pl	acebo			
Atmaca 2002	9/14	4/13		2.09 [0.85, 5.16]
Subtotal (95% CI)	14	13		2.09 [0.85, 5.16]
Total events: 9 (Augmentati	ion), 4 (non-augmentation)		-	
Test for heterogeneity: not	applicable			
Test for overall effect: Z = 1	1.60 (P = 0.11)			
02 SRI+Risperidone v SRI+F	Placebo			
Hollander 2003e	4/10	2/6		1.20 [0.31, 4.69]
McDougle 2000a	18/20	15/16	<u> </u>	0.96 [0.79, 1.16]
Subtotal (95% CI)	30	22	•	0.99 [0.77, 1.28]
Total events: 22 (Augments	ation), 17 (non-augmentation)			
Test for heterogeneity: Chi ²	² = 0.18, df = 1 (P = 0.67), l ² = 09	6		
Test for overall effect: Z = I	0.07 (P = 0.95)			
Total (95% CI)	44	35	•	1.19 [0.89, 1.58]
Total events: 31 (Augments	ation), 21 (non-augmentation)			, ,
Test for heterogeneity: Chi ²	$^{2} = 6.13$, df = 2 (P = 0.05), $l^{2} = 67$.3%		
Test for overall effect: Z = 1				
			0.1 0.2 0.5 1 2 5	10
		F	avours augmentation Favours non-au	g

Review: OCD: augmentation strategies 12 All antipsychotics v placebo Comparison: Outcome: 02 Leaving the study early

Study or sub-category	Augmentation n/N	non-augmentation n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 FLX+Olanzapine v FLX+Plac	cebo			
Shapira 2004	5/22	2/22		2.50 [0.54, 11.54]
Subtotal (95% CI)	22	22		2.50 [0.54, 11.54]
otal events: 5 (Augmentation)	, 2 (non-augmentation)			,
est for heterogeneity: not app	licable			
est for overall effect: Z = 1.17				
)2 SRI+Quetiapine v SRI+Place	bo			
Atmaca 2002	0/14	0/13		Not estimable
Subtotal (95% CI)	0	0		Not estimable
「otal events: 0 (Augmentation)	ı, 0 (non-augmentation)			
Test for heterogeneity: not app	olicable			
est for overall effect: not appl	licable			
03 SRI+Risperidone v SRI+Plac	ebo			
Hollander 2003e	1/10	2/6		0.30 [0.03, 2.65]
McDougle 2000a	2/20	1/16	- -	1.60 [0.16, 16.10]
Subtotal (95% CI)	30	22	-	0.70 [0.16, 3.01]
otal events: 3 (Augmentation)	ı, 3 (non-augmentation)			
'est for heterogeneity: Chi² = 1	l.07, df = 1 (P = 0.30), l² = 6.9%			
est for overall effect: Z = 0.48	3 (P = 0.63)			
otal (95% CI)	66	57	-	1.34 [0.49, 3.67]
「otal events: 8 (Augmentation)	ı, 5 (non-augmentation)		[-	
est for heterogeneity: Chi² = 2	2.48, df = 2 (P = 0.29), l ² = 19.3%			
Test for overall effect: $Z = 0.57$	7 (P = 0.57)			
			0.01 0.1 1 10	100
			Favours augmentation Favours non-au	ua

Review: OCD: augmentation strategies Comparison: 12 All antipsychotics v placebo

Outcome: 03 Leaving the study early due to adverse effects

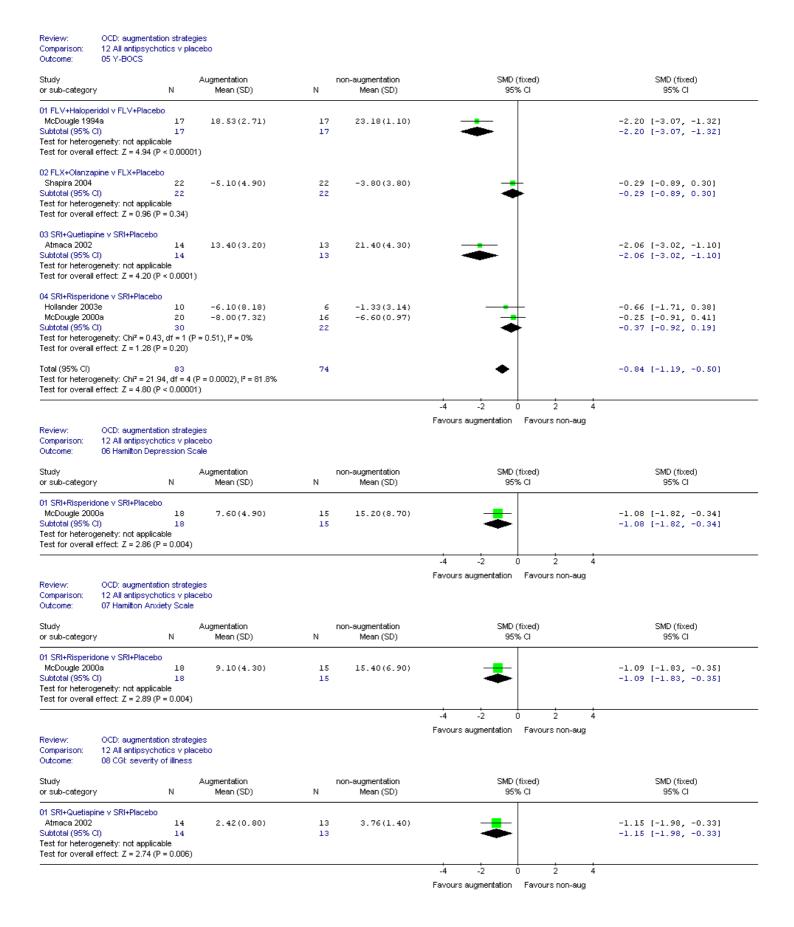


Review: OCD: augmentation strategies
Comparison: 12 All antipsychotics v placebo

Outcome: 04 Non-responders

Study or sub-category	Augmentation n/N	non-augmentation n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl		
01 FLV+Haloperidol v FLV+Placebo						
McDougle 1994a	10/17	17/17		0.59 [0.40, 0.88]		
Subtotal (95% CI)	17	17	-	0.59 [0.40, 0.88]		
Total events: 10 (Augmentation), 17	, -					
Test for heterogeneity: not applicab						
Test for overall effect: Z = 2.61 (P =	= 0.009)					
02 FLX+Olanzapine v FLX+Placebo						
Shapira 2004	13/22	13/22		1.00 [0.61, 1.64]		
Subtotal (95% CI)	22	22	-	1.00 [0.61, 1.64]		
Total events: 13 (Augmentation), 13	(non-augmentation)		Ī			
Test for heterogeneity: not applicab						
Test for overall effect: Z = 0.00 (P =	= 1.00)					
03 SRI+Quetiapine v SRI+Placebo						
Atmaca 2002	4/14	13/13 —		0.29 [0.12, 0.65]		
Subtotal (95% CI)	14	13 -		0.29 [0.12, 0.65]		
Total events: 4 (Augmentation), 13						
Test for heterogeneity: not applicab						
Test for overall effect: Z = 2.96 (P =	= 0.003)					
04 SRI+Risperidone v SRI+Placebo						
Hollander 2003e	6/10	6/6		0.60 [0.36, 1.00]		
McDougle 2000a	16/20	16/16	-	0.80 [0.64, 1.00]		
Subtotal (95% CI)	30	22	◆	0.74 [0.60, 0.91]		
Total events: 22 (Augmentation), 22						
Test for heterogeneity: Chi ² = 1.14,		.3%				
Test for overall effect: Z = 2.81 (P =	= 0.005)					
Total (95% CI)	83	74	•	0.66 [0.54, 0.81]		
Total events: 49 (Augmentation), 65	(non-augmentation)		, l	•		
Test for heterogeneity: Chi ² = 9.98,	$df = 4 (P = 0.04), I^2 = 59$.9%				
Test for overall effect: Z = 4.05 (P <	< 0.0001)					
		0.1	0.2 0.5 1 2 5	10		

Favours augmentation Favours non-aug



SSRIs (OCD, children)

Review: OCD children: SSRIs
Comparison: 01 SSRI vs placebo (acute phase)
Outcome: 01 Adverse events

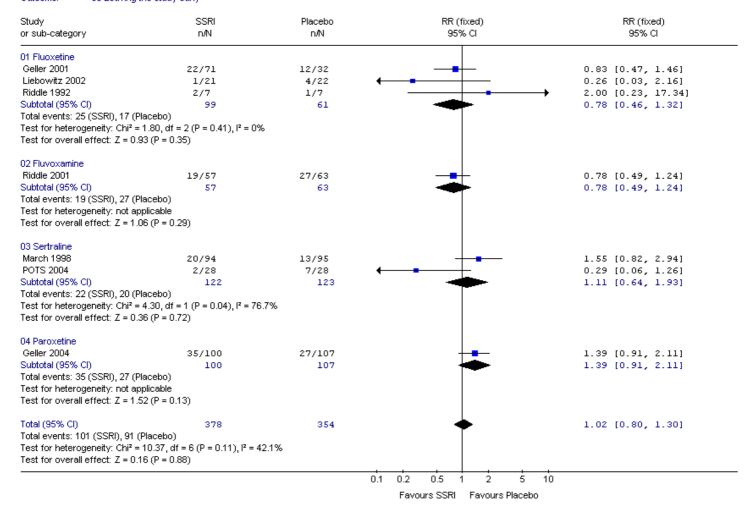
Study or sub-category	SSRI n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Fluoxetine				
Geller 2001	53/71	27/32		0.88 [0.72, 1.08]
Liebowitz 2002	21/21	19/22		1.16 [0.98, 1.37]
Subtotal (95% CI)	92	54	•	0.98 [0.85, 1.12]
Total events: 74 (SSRI), 46 (F	Placebo)		Ţ	
Test for heterogeneity: Chi ² = Test for overall effect: $Z = 0.3$	= 4.99, df = 1 (P = 0.03), l² = 80 34 (P = 0.73)).0%		
02 Fluvoxamine	40.458	40.450		
Riddle 2001	48/57	48/63		1.11 [0.93, 1.32]
Subtotal (95% CI) Total events: 48 (SSRI), 48 (F Test for heterogeneity: not ap Test for overall effect: Z = 1.	oplicable	63		1.11 [0.93, 1.32]
03 Paroxetine				
Geller 2004	83/98	77/105		1.15 [1.00, 1.33]
Subtotal (95% CI) Total events: 83 (SSRI), 77 (F Test for heterogeneity: not ap Test for overall effect: Z = 1.9	oplicable	105		1.15 [1.00, 1.33]
	= 5.30, df = 3 (P = 0.15), l ² = 43	222	•	1.09 [0.99, 1.19]
Test for overall effect: Z = 1.3	80 (P = 0.07)			
		0.5	0.7 1 1.5	2
			Favours SSRI Favours Placeb	0

Review: OCD children: SSRIs
Comparison: 01 SSRI vs placebo (acute phase) Outcome: 02 Serious adverse events

Study or sub-category	SSRI n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Paroxetine				
Geller 2004	3/98	1/105	- - - - - - - - - - 	- 3.21 [0.34, 30.39]
Subtotal (95% CI)	98	105		3.21 [0.34, 30.39]
Total events: 3 (SSRI), 1 (Placebo)				
Test for heterogeneity: not applicable				
Test for overall effect: $Z = 1.02$ (P = 0.31)			
Total (95% CI)	98	105		- 3.21 [0.34, 30.39]
Total events: 3 (SSRI), 1 (Placebo)			- - -	
Test for heterogeneity: not applicable				
Test for overall effect: $Z = 1.02$ (P = 0.31)			
		0.01	0.1 1 10	100
			Favours SSRI Favours Plac	cebo

Review: OCD children: SSRIs

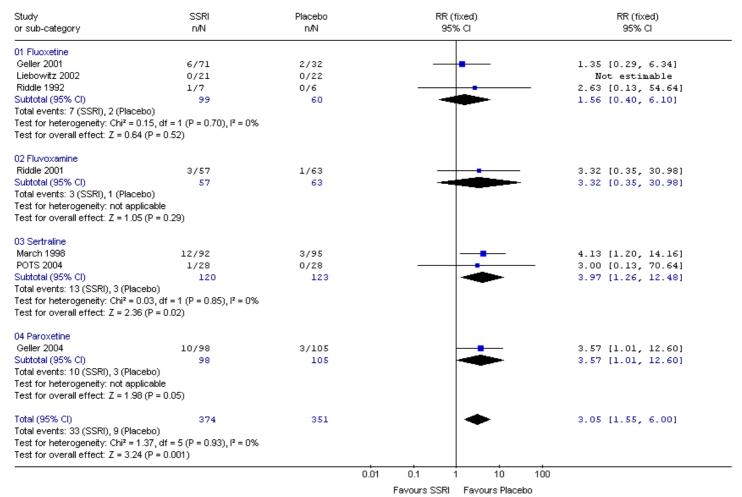
Comparison: 01 SSRI vs placebo (acute phase)
Outcome: 03 Leaving the study early



Review: OCD children: SSRIs

Comparison: 01 SSRI vs placebo (acute phase)

Outcome: 04 Leaving the study early due to adverse events



Review: OCD children: SSRIs

Comparison: 01 SSRI vs placebo (acute phase)
Outcome: 05 Non-responders (25% CY-BOCS)

Study or sub-category	SSRI n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Fluvoxamine				
Riddle 2001	33/57	46/63		0.79 [0.61, 1.04]
Subtotal (95% CI)	57	63	•	0.79 [0.61, 1.04]
Total events: 33 (SSRI), 46 (Place	ebo)		-	•
Test for heterogeneity; not applica	able			
Test for overall effect: $Z = 1.70$ (Fig. 1)	P = 0.09)			
02 Paroxetine				
Geller 2004	39/100	65/107		0.64 [0.48, 0.86]
Subtotal (95% CI)	100	107	◆	0.64 [0.48, 0.86]
Total events: 39 (SSRI), 65 (Place	ebo)			
Test for heterogeneity: not applicate	able			
Test for overall effect: Z = 3.01 (F	P = 0.003)			
Total (95% CI)	157	170	•	0.70 [0.58, 0.86]
Total events: 72 (SSRI), 111 (Plac	cebo)		-	
Test for heterogeneity: Chi ² = 1.1	5, df = 1 (P = 0.28), l² = 13	.2%		
Test for overall effect: $Z = 3.44$ (i	P = 0.0006)			
		0.1	0.2 0.5 1 2 5	10
			Favours SSRI Favours Placel	bo

Review: OCD children: SSRIs Comparison: 01 SSRI vs placebo (acute phase) Outcome: 06 Non-responders (40% CY-BOCS) SSRI Placebo RR (fixed) RR (fixed) Study or sub-category nN95% CI 95% CI 01 Fluoxetine Geller 2001 36/71 24/32 0.68 [0.50, 0.92] Subtotal (95% CI) 71 32 0.68 [0.50, 0.92] Total events: 36 (SSRI), 24 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 2.52 (P = 0.01) 0.1 0.2 0.5 ż 5 Favours SSRI Favours Placebo Review: OCD children: SSRIs Comparison: 01 SSRI vs placebo (acute phase) Outcome: 07 Non-responders (all) Study SSRI Placebo RR (fixed) RR (fixed) or sub-category n/Ν n/N 95% CI 95% CI 01 Fluoxetine Geller 2001 36/71 24/32 0.68 [0.50, 0.92] Subtotal (95% CI) 0.68 [0.50, 0.92] 71 32 Total events: 36 (SSRI), 24 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 2.52 (P = 0.01)02 Fluvoxamine 0.79 [0.61, 1.04] Riddle 2001 33/57 46/63 Subtotal (95% CI) 0.79 [0.61, 1.04] 57 63 Total events: 33 (SSRI), 46 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.70 (P = 0.09) 03 Paroxetine Geller 2004 0.64 [0.48, 0.86] 39/100 65/107 Subtotal (95% CI) 100 107 0.64 [0.48, 0.86] Total events: 39 (SSRI), 65 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 3.01 (P = 0.003) 0.70 [0.59, 0.83] Total (95% CI) 228 202 Total events: 108 (SSRI), 135 (Placebo) Test for heterogeneity: $Chi^2 = 1.24$, df = 2 (P = 0.54), $I^2 = 0\%$ Test for overall effect: Z = 4.18 (P < 0.0001) 0.5 0.2 Favours SSRI Favours placebo OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome: 08 Non-responders (CGI Global Improvement) RR (fixed) Study SSRI Placebo RR (fixed) or sub-category n/N n/N 95% CI 95% CI

Management of OCD: full guideline DRAFT (May 2005)

55/100

72/107

0.1 0.2

0.5

Favours SSRI

0.82 [0.66, 1.02]

0.82 [0.66, 1.02]

5

Favours Placebo

10

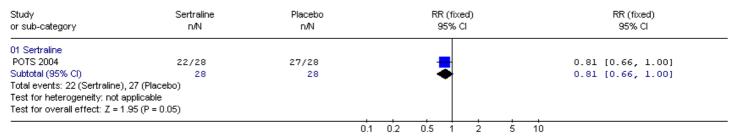
01 Paroxetine Geller 2004

Subtotal (95% CI)

Total events: 55 (SSRI), 72 (Placebo)
Test for heterogeneity: not applicable
Test for overall effect: Z = 1.79 (P = 0.07)

OCD children: SSRIs Review:

01 SSRI vs placebo (acute phase) Comparison: 09 Remission (CY-BOCS < 11) Outcome:



Favours Sertraline Favours Placebo

OCD children: SSRIs Review:

Comparison: 01 SSRI vs placebo (acute phase)

Outcome:

Study or sub-category	N	SSRI Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
Geller 2001	71	-9.50(9.20)	32	-5.20(7.40)		-0.49 [-0.91, -0.07]
Liebowitz 2002	21	14.71(8.73)	22	18.55(11.44)		-0.37 [-0.97, 0.23]
Riddle 1992	7	13.60(5.70)	-6	14.80(7.00)		-0.18 [-1.27, 0.92]
Subtotal (95% CI)	99	10.00(0.70)	60	14.00(7.007	`	-0.43 [-0.76, -0.10]
Test for heterogeneity: Chi² =		- 0.95) 12 - 0%	- 00			-0.43 [-0.76, -0.10]
rest for neterogeneity. Cril = Test for overall effect: Z = 2.5		-0.03),1 -0.00				
100110101010110111101111111111111111111						
02 Fluvoxamine						
Riddle 2001	57	18.20(8.60)	63	20.90(8.50)	- •	-0.31 [-0.67, 0.05]
Subtotal (95% Cl)	57		63		-	-0.31 [-0.67, 0.05]
Test for heterogeneity: not app						
Test for overall effect: $Z = 1.7$	1 (P = 0.09)					
03 Sertraline						
March 1998	92	-6.80(8.35)	95	-3.40(8.00)		-0.41 [-0.70, -0.12]
POTS 2004	28	16.50(9.10)	28	21.50(5.40)		-0.66 [-1.20, -0.12]
Subtotal (95% CI)	120		123			-0.47 [-0.72, -0.21]
Test for heterogeneity: Chi ² =	0.61, df = 1 (P	= 0.43), I ² = 0%			_	
Test for overall effect: Z = 3.6	0 (P = 0.0003)					
04 Paroxetine						
Geller 2004	94	-8.78(7.65)	102	-5.34(7.78)		-0.44 [-0.73, -0.16]
Subtotal (95% CI)	94		102			-0.44 [-0.73, -0.16]
Test for heterogeneity: not ap						
Test for overall effect: Z = 3.0						
Total (95% CI)	370		348			-0.43 [-0.58, -0.28]
Test for heterogeneity: Chi ² =		= 0.96) I ² = 0%	040		_	0.10 (0.00, 0.20)
Test for overall effect: Z = 5.5						
					-1 -0.5 0 0.5	i
					Favours SSRI Favours Pla	cebo

Review: OCD children: SSRIs

Comparison: Outcome: 01 SSRI vs placebo (acute phase) 11 NIMH-OC

Study or sub-category	N	SSRI Mean (SD)	N	Placebo Mean (SD)		SMD (fixe 95% CI	d)	SMD (fixed) 95% Cl
01 Fluoxetine								
Geller 2001	71	-3.10(3.00)	32	-1.30(2.20)	←	<u> </u>		-0.64 [-1.07, -0.22]
Liebowitz 2002	21	6.24(2.45)	22	7.18(2.75)		-	_	-0.35 [-0.96, 0.25]
Subtotal (95% CI)	92		54					-0.55 [-0.89, -0.20]
Test for heterogeneity: Chi ² =	0.59, df = 1 (P	= 0.44), I ² = 0%				_		
Test for overall effect: $Z = 3.0$	07 (P = 0.002)							
02 Fluvoxamine								
Riddle 2001	57	7.50(3.20)	63	8.30(2.60)				-0.27 [-0.63, 0.09]
Subtotal (95% CI)	57		63					-0.27 [-0.63, 0.09]
Test for heterogeneity: not app								
Test for overall effect: $Z = 1.4$	19 (P = 0.14)							
03 Sertraline								
March 1998	92	-2.20(2.78)	95	-1.30(2.63)				-0.33 [-0.62, -0.04]
Subtotal (95% CI)	92		95					-0.33 [-0.62, -0.04]
Test for heterogeneity: not app	plicable							
Test for overall effect: $Z = 2.2$	25 (P = 0.02)							
Total (95% CI)	241		212			•		-0.38 [-0.57, -0.19]
Test for heterogeneity: Chi2 =	1.91, df = 3 (P	= 0.59), I ² = 0%				-		
Test for overall effect: $Z = 3.9$	93 (P < 0.0001)							
					-1	-0.5	0.5	1
					Fa	avours SSRI Fa	vours Placebo	

OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome 12 Child Obsessive Compulsive Impact Scale (Parent Version) SMD (fixed) SMD (fixed) Study Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 21 38.44(30.43) 22 53.95(41.97) -0.41 [-1.02, 0.19] Subtotal (95% CI) 21 22 -0.41 [-1.02, 0.19] Test for heterogeneity: not applicable Test for overall effect: Z = 1.34 (P = 0.18) ò Favours SSRI Favours Placebo OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome 13 Leyton Obsessional Inventory (Child Version): interference SMD (fixed) Study SSRI SMD (fixed) Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Riddle 1992 7 20.60(19.40) 6 21.00(19.30) -0.02 [-1.11, 1.07] Subtotal (95% CI) -0.02 [-1.11, 1.07] Test for heterogeneity: not applicable Test for overall effect: Z = 0.03 (P = 0.97) Favours SSRI Favours Placebo Review: OCD children: SSRIs 01 SSRI vs placebo (acute phase) Comparison: Outcome: 14 Leyton Obsessional Inventory (Child Version): resistance Study SSRI Placebo SMD (fixed) SMD (fixed) Ν Mean (SD) Ν or sub-category Mean (SD) 95% CI 95% CI 01 Fluoxetine Riddle 1992 19.60(14.40) 22.00(23.10) -0.12 [-1.21, 0.97] Subtotal (95% CI) -0.12 [-1.21, 0.97] Test for heterogeneity; not applicable Test for overall effect: Z = 0.21 (P = 0.83) -4 -2 ń Favours SSRI Favours Placebo OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome 15 Leyton Obsessional Inventory (Child Version): symptoms SMD (fixed) Study SSRI Placebo SMD (fixed) or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Riddle 1992 0.23 [-0.87, 1.32] 16.30(8.40) 6 13.80(12.10) Subtotal (95% CI) 0.23 [-0.87, 1.32] Test for heterogeneity: not applicable Test for overall effect: Z = 0.41 (P = 0.68) Favours SSRI Favours Placebo OCD children; SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome 16 Children's Depression Rating Scale (Revised Version) SSRI SMD (fixed) SMD (fixed) Study Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluvoxamine Riddle 2001 57 27.40(8.00) 63 27.10(8.00) 0.04 [-0.32, 0.40] 0.04 [-0.32, 0.40] Subtotal (95% CI) 57 63 Test for heterogeneity: not applicable Test for overall effect: Z = 0.20 (P = 0.84) -0.5 0.5 Ó Favours SSRI Favours Placebo Review: OCD children: SSRIs Comparison: 01 SSRI vs placebo (acute phase) Outcome 17 Hamilton Rating Scale for Depression SSRI Placebo SMD (fixed) SMD (fixed) Study or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 21 9.14(6.56) 7.09(7.39) 0.29 [-0.31, 0.89] 22 Subtotal (95% CI) 21 22 0.29 [-0.31, 0.89] Test for heterogeneity; not applicable Test for overall effect: Z = 0.94 (P = 0.35) -0.5 Ó 0.5

Favours SSRI Favours Placebo

OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome 18 Depression (CDRS; HRSD) SMD (fixed) SMD (fixed) Study Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 21 9.14(6.56) 22 7.09(7.39) 0.29 [-0.31, 0.89] Subtotal (95% CI) 21 22 0.29 [-0.31, 0.89] Test for heterogeneity: not applicable Test for overall effect: Z = 0.94 (P = 0.35) Riddle 2001 57 27.40(8.00) 63 27.10(8.00) 0.04 [-0.32, 0.40] Subtotal (95% CI) 57 63 0.04 [-0.32, 0.40] Test for heterogeneity; not applicable Test for overall effect: Z = 0.20 (P = 0.84) 0.10 [-0.20, 0.41] Total (95% CI) 85 Test for heterogeneity: Chi² = 0.49, df = 1 (P = 0.48), l² = 0% Test for overall effect: Z = 0.65 (P = 0.51) -0.5 ń 0.5 Favours SSRI Favours placebo Review: OCD children: SSRIs Comparison: 01 SSRI vs placebo (acute phase) Outcome 19 Multidimensional Anxiety Scale for Children SMD (fixed) SMD (fixed) Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Geller 2001 71 -9.00(18.60) 32 -4.50(16.50) -0.25 [-0.67, 0.17] Subtotal (95% CI) 32 -0.25 [-0.67, 0.17] Test for heterogeneity: not applicable Test for overall effect: Z = 1.16 (P = 0.24) -0.5 Ö 0.5 Favours SSRI Favours Placebo OCD children: SSRIs Review: Comparison: 01 SSRI vs placebo (acute phase) Outcome: 20 Revised Children's Manifest Anxiety Scale SMD (fixed) SSRI SMD (fixed) Study Placebo or sub-category Ν Mean (SD) N Mean (SD) 95% CI 95% CI 01 Fluoxetine Riddle 1992 45.70(13.30) 40.20(16.90) 0.34 [-0.86, 1.53] Subtotal (95% CI) 0.34 [-0.86, 1.53] Test for heterogeneity: not applicable Test for overall effect: Z = 0.55 (P = 0.58) -4 0 Favours SSRI Favours Placebo Review: OCD children: SSRIs 01 SSRI vs placebo (acute phase) Comparison: 21 Anxiety (MASC; RCMAS) Outcome Study SSRI Placebo SMD (fixed) SMD (fixed) Ν Mean (SD) Ν Mean (SD) or sub-category 95% CI 95% CI 01 Fluoxetine Geller 2001 71 -9.00(18.60) 32 -4.50(16.50) -0.25 [-0.67, 0.17] 0.34 [-0.86, 1.53] -0.19 [-0.58, 0.21] Riddle 1992 6 45.70(13.30) 5 40.20(16.90) Subtotal (95% CI) 37 77 Test for heterogeneity: Chi² = 0.81, df = 1 (P = 0.37), l² = 0% Test for overall effect: Z = 0.92 (P = 0.36) -2 0 Favours SSRI Favours placebo Review: OCD children: SSRIs Comparison: 01 SSRI vs placebo (acute phase) 22 Children's Global Assessment Scale Outcome Study SSRI Placebo SMD (fixed) SMD (fixed) or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Riddle 1992 70.40(13.20) 62.00(8.40) 0.69 [-0.44, 1.83] Subtotal (95% CI) 0.69 [-0.44, 1.83] Test for heterogeneity: not applicable Test for overall effect: Z = 1.20 (P = 0.23)

-2

Ò Favours SSRI Favours Placebo

Review: OCD children: SSRIs

Comparison: 01 SSRI vs placebo (acute phase) Outcome: 23 Systolic blood pressure - supine

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Sertraline v Placebo (at 1	week)					
March 1998	92	108.50(12.50)	95	107.60(10.60)	<u>=</u>	0.08 [-0.21, 0.36]
Subtotal (95% Cl)	92		95		→	0.08 [-0.21, 0.36]
Test for heterogeneity: not a	pplicable					
Test for overall effect: $Z = 0$	53 (P = 0.60)					
02 Sertraline v Placebo (at 4	weeks)					
March 1998	92	108.10(12.10)	95	107.00(11.00)	iii	0.09 [-0.19, 0.38]
Subtotal (95% CI)	92		95		T	0.09 [-0.19, 0.38]
Fest for heterogeneity: not a	pplicable				ľ	
Fest for overall effect: $Z = 0$	65 (P = 0.52)					
03 Sertraline v Placebo (at 1:	2 weeks)					
March 1998	92	107.00(11.00)	95	107.40(10.60)	<u>=</u>	-0.04 [-0.32, 0.25]
Subtotal (95% CI)	92		95			-0.04 [-0.32, 0.25]
Test for heterogeneity: not a	pplicable					
Test for overall effect: $Z = 0$.	25 (P = 0.80)					

Review:

OCD children: SSRIs 01 SSRI vs placebo (acute phase) Comparison: Outcome: 24 Systolic blood pressure - standing

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% CI	SMD (fixed) 95% CI
01 Sertraline v Placebo (at 1 v	week)					
March 1998	92	108.40(11.00)	95	107.40(10.90)	<u> </u>	0.09 [-0.20, 0.38]
Subtotal (95% CI)	92		95		<u>∓</u>	0.09 [-0.20, 0.38]
Test for heterogeneity: not ap	plicable				ſ	
Test for overall effect: $Z = 0.6$	62 (P = 0.53)					
02 Sertraline v Placebo (at 4 v	weeks)					
March 1998	92	106.20(11.40)	95	107.20(11.70)	<u> </u>	-0.09 [-0.37, 0.20]
Subtotal (95% CI)	92		95			-0.09 [-0.37, 0.20]
Test for heterogeneity: not ap	plicable					
Test for overall effect: Z = 0.5	59 (P = 0.56)					
03 Sertraline v Placebo (at 12	weeks)					
March 1998	92	105.80(10.80)	95	107.10(10.90)	<u> </u>	-0.12 [-0.41, 0.17]
Subtotal (95% CI)	92	•	95		7	-0.12 [-0.41, 0.17]
Test for heterogeneity: not ap	plicable]	•
Test for overall effect: $Z = 0.8$						

Favours sertraline Favours placebo

Favours sertraline Favours placebo

Favours sertraline Favours placebo

OCD children: SSRIs Comparison: Outcome: 01 SSRI vs placebo (acute phase) 25 Diastolic blood pressure - supine

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% Cl
01 Sertraline v Placebo (at 1	week)					
March 1998	92	64.80(8.50)	95	64.40(10.20)	in the second se	0.04 [-0.24, 0.33]
Subtotal (95% CI)	92		95		→	0.04 [-0.24, 0.33]
Test for heterogeneity: not ap	plicable				ſ	
Test for overall effect: $Z = 0$.	29 (P = 0.77)					
02 Sertraline v Placebo (at 4	weeks)					
March 1998	92	63.00(8.70)	95	63.80(9.40)	in the second se	-0.09 [-0.37, 0.20]
Subtotal (95% CI)	92		95		<u>▼</u>	-0.09 [-0.37, 0.20]
Test for heterogeneity: not ap	plicable]	
Test for overall effect: $Z = 0$.	60 (P = 0.55)					
03 Sertraline v Placebo (at 12	! weeks)					
March 1998	92	63.70(10.90)	95	64.50(8.80)	e e e e e e e e e e e e e e e e e e e	-0.08 [-0.37, 0.21]
Subtotal (95% CI)	92		95		<u>▼</u>	-0.08 [-0.37, 0.21]
Test for heterogeneity: not ap	plicable				1	•
Test for overall effect: $Z = 0$.						

Review: OCD children: SSRIs

01 SSRI vs placebo (acute phase) 26 Diastolic blood pressure - standing Comparison: Outcome:

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% CI
01 Sertraline v Placebo (at 1 week)						
March 1998	92	67.60(9.40)	95	67.70(10.60)	<u> </u>	-0.01 [-0.30, 0.28]
Subtotal (95% CI)	92		95		∓	-0.01 [-0.30, 0.28]
Test for heterogeneity: not applicable	е				1	
Test for overall effect: Z = 0.07 (P =	0.95)					
02 Sertraline v Placebo (at 4 weeks))					
March 1998	92	67.20(10.60)	95	67.40(9.80)	<u> </u>	-0.02 [-0.31, 0.27]
Subtotal (95% CI)	92		95			-0.02 [-0.31, 0.27]
Test for heterogeneity: not applicable	е					·
Test for overall effect: $Z = 0.13$ (P =	0.89)					
03 Sertraline v Placebo (at 12 weeks	s)					
March 1998	92	67.10(9.20)	95	67.80(8.90)	<u> </u>	-0.08 [-0.36, 0.21]
Subtotal (95% CI)	92		95			-0.08 [-0.36, 0.21]
Test for heterogeneity: not applicable	е				1	
Test for overall effect: Z = 0.53 (P =	0.60)					
				-10	-5 0 5	10

Favours sertraline Favours placebo

Favours sertraline Favours placebo

Review:

Review: OCD children: SSRIs
Comparison: 01 SSRI vs placebo (acute phase)
Outcome: 27 Heart rate - supine

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)	SMD (fixed) 95% Cl	SMD (fixed) 95% CI
01 Sertraline v Placebo (at 1 week)						
March 1998	92	75.40(12.70)	95	77.80(13.00)	<u>=</u>	-0.19 [-0.47, 0.10]
Subtotal (95% CI)	92		95		₹	-0.19 [-0.47, 0.10]
Test for heterogeneity; not applicable	е]	
Test for overall effect: Z = 1.27 (P =	0.20)					
02 Sertraline v Placebo (at 4 weeks))					
March 1998	92	76.90(11.50)	95	78.10(11.80)	<u>=</u>	-0.10 [-0.39, 0.18]
Subtotal (95% CI)	92		95			-0.10 [-0.39, 0.18]
Test for heterogeneity; not applicable	е					
Test for overall effect: $Z = 0.70$ (P =	0.48)					
03 Sertraline v Placebo (at 12 weeks	s)					
March 1998	92	77.30(9.80)	95	78.40(13.00)	<u> </u>	-0.09 [-0.38, 0.19]
Subtotal (95% CI)	92		95			-0.09 [-0.38, 0.19]
Test for heterogeneity: not applicable	е					
Test for overall effect: Z = 0.65 (P =						
				-10	-5 0 5	10

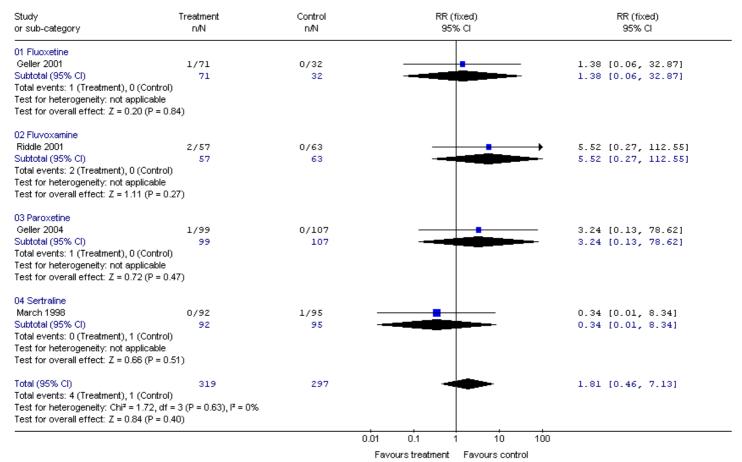
OCD children: SSRIs Comparison: Outcome: 01 SSRI vs placebo (acute phase) 28 Heart rate - standing

Study or sub-category	N	Sertraline Mean (SD)	N	Placebo Mean (SD)		SMD (fixed) 95% CI		SMD (fixed) 95% Cl
01 Sertraline v Placebo (at 1 °	week)							
March 1998	92	81.40(13.90)	95	86.10(13.30)				-0.34 [-0.63, -0.06]
Subtotal (95% CI)	92		95			•		-0.34 [-0.63, -0.06]
Test for heterogeneity: not ap	pplicable					1		
Test for overall effect: $Z = 2.3$	34 (P = 0.02)							
02 Sertraline v Placebo (at 4 °	weeks)							
March 1998	92	83.90(13.80)	95	84.50(12.00)				-0.05 [-0.33, 0.24]
Subtotal (95% CI)	92		95			•		-0.05 [-0.33, 0.24]
Test for heterogeneity: not ap	pplicable							
Test for overall effect: $Z = 0.3$	32 (P = 0.75)							
03 Sertraline v Placebo (at 12	2 weeks)							
March 1998	92	85.10(12.10)	95	86.20(15.30)				-0.08 [-0.37, 0.21]
Subtotal (95% CI)	92		95			•		-0.08 [-0.37, 0.21]
Test for heterogeneity: not ap	pplicable							
Test for overall effect: $Z = 0.3$	54 (P = 0.59)							
					-105	5 0	5	10
					Favourss	ertraline Favo	urs placebo	i

Review: OCD children: SSRIs

Comparison: 01 SSRI vs placebo (acute phase)

Outcome: 29 Suicidality (Columbia reclassification - "Outcome 3: definitive suicidal behaviour/ ideation")



Review: OCD children: SSRIs

Comparison: 02 SSRI vs placebo (continuation phase)

Outcome: 01 Adverse events

Subtotal (95% CI) 11 7 - 4.67 Total events: 3 (SSRI), 0 (Placebo)	
Subtotal (95% CI) 11 7	
Total events: 3 (SSRI), 0 (Placebo)	[0.28, 78.68]
, ,, ,	[0.28, 78.68]
Test for heterogeneity: not applicable	
Test for overall effect: Z = 1.07 (P = 0.29)	

Favours SSRI

Favours Placebo

Review: OCD children: SSRIs

Comparison: 02 SSRI vs placebo (continuation phase)

Outcome: 02 Leaving study early

Study or sub-category	SSRI n/N	Placebo n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
01 Fluoxetine Liebowitz 2002	1/12	0/7		- 1.85 [0.09, 40.05]
Subtotal (95% CI)	1/12	0/7		- 1.85 [0.09, 40.05]
Test for heterogeneity: not ap Test for overall effect: Z = 0.3	•			
		0.	01 0.1 1 10	100
			Favours SSRI Favours Place	bo

OCD children: SSRIs Review: Comparison: 02 SSRI vs placebo (continuation phase) 03 CY-BOCS Outcome SMD (fixed) SMD (fixed) Study Placebo or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 11 5.64(5.41) 11.43(11.12) -0.69 [-1.67, 0.30] Subtotal (95% CI) 11 -0.69 [-1.67, 0.30] Test for heterogeneity: not applicable Test for overall effect; Z = 1.37 (P = 0.17) -10 ò 5 10 Favours SSRI Favours Placebo OCD children: SSRIs Review: Comparison: 02 SSRI vs placebo (continuation phase) Outcome 04 NIMH-OC SMD (fixed) SMD (fixed) Study SSRI Placebo or sub-category N Mean (SD) Mean (SD) 95% CI 95% CI 01 Fluoxetine -0.69 [-1.67, 0.29] -0.69 [-1.67, 0.29] Liebowitz 2002 11 3.73(1.74) 7 5.43(3.10) Subtotal (95% CI) 11 Test for heterogeneity: not applicable Test for overall effect: Z = 1.38 (P = 0.17) -10 -5 0 Favours SSRI Favours Placebo Review: OCD children: SSRIs 02 SSRI vs placebo (continuation phase) Comparison: Outcome: 05 Child Obsessive Compulsive Impact Scale (Parent version) Study SSRI Placebo SMD (fixed) SMD (fixed) Ν Mean (SD) Ν Mean (SD) or sub-category 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 11 23.00(20.57) 49.29(32.70) -0.97 [-1.98, 0.04] Subtotal (95% CI) 11 -0.97 [-1.98, 0.04] Test for heterogeneity; not applicable Test for overall effect: Z = 1.88 (P = 0.06) -100 -50 Ó 50 100 Favours SSRI Favours Placebo OCD children: SSRIs Comparison: 02 SSRI vs placebo (continuation phase) Outcome 06 Hamilton Rating Scale for Depression SMD (fixed) SMD (fixed) Study SSRI Placebo or sub-category Ν Mean (SD) Ν Mean (SD) 95% CI 95% CI 01 Fluoxetine Liebowitz 2002 -0.41 [-1.37, 0.55] 11 4.64(6.14) 7.14(5.24) Subtotal (95% CI) 11 -0.41 [-1.37, 0.55] Test for heterogeneity: not applicable Test for overall effect: Z = 0.84 (P = 0.40) -10 10 Favours SSRI Favours Placebo Review: OCD children: SSRIs Comparison: 03 SSRIs vs placebo (discontinuation) Outcome: 01 Adverse events RR (fixed) RR (fixed) Study Paroxetine Placebo 95% CI 95% CI or sub-category n/N n/N Carpenter 1999 73/95 80/98 0.94 [0.81, 1.09] Total (95% CI) 98 0.94 [0.81, 1.09] 95 Total events: 73 (Paroxetine), 80 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.82 (P = 0.41)

0.5

0.7

Favours Paroxetine

1.5

Favours Placebo

ż

Review: OCD children: SSRIs 03 SSRIs vs placebo (discontinuation) Comparison: Outcome: 02 Serious adverse events Paroxetine Placebo RR (fixed) RR (fixed) Study 95% CI 95% CI or sub-category n/N n/N Carpenter 1999 2/95 2/98 1.03 [0.15, 7.18] Total (95% CI) 95 98 1.03 [0.15, 7.18] Total events: 2 (Paroxetine), 2 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.03 (P = 0.97) 0.1 0.2 0.5 5 10 Favours Paroxetine Favours Placebo Review: OCD children: SSRIs Comparison: 03 SSRIs vs placebo (discontinuation) 03 Leaving the study early Outcome: Study RR (fixed) RR (fixed) Paroxetine Placebo or sub-category n/N n/N 95% CI 95% CI Carpenter 1999 54/96 65/98 0.85 [0.68, 1.06] Total (95% CI) 96 98 0.85 [0.68, 1.06] Total events: 54 (Paroxetine), 65 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 1.43 (P = 0.15) 0.5 0.7 1.5 ż Favours Paroxetine Favours Placebo OCD children: SSRIs Review: 03 SSRIs vs placebo (discontinuation) Comparison: Outcome: 04 Leaving the study early due to adverse events RR (fixed) RR (fixed) Study Paroxetine Placebo n/N 95% CI 95% CI or sub-category n/N Carpenter 1999 8/95 11/98 0.75 [0.32, 1.78] Total (95% CI) 98 0.75 [0.32, 1.78] Total events: 8 (Paroxetine), 11 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 0.65 (P = 0.52) 0.1 0.2 0.5 2 5 10 Favours Paroxetine Favours Placebo OCD children: SSRIs Review: Comparison: 03 SSRIs vs placebo (discontinuation) Outcome: 05 Non-responders (25% CY-BOCS) Study Paroxetine Placebo RR (fixed) RR (fixed) or sub-category n/Ν n/Ν 95% CI 95% CI Carpenter 1999 72/96 85/98 0.86 [0.75, 0.99] Total (95% CI) 96 98 0.86 [0.75, 0.99] Total events: 72 (Paroxetine), 85 (Placebo) Test for heterogeneity: not applicable Test for overall effect: Z = 2.05 (P = 0.04) Favours Paroxetine Favours Placebo OCD children: SSRIs Review: Comparison: 03 SSRIs vs placebo (discontinuation) Outcome: 06 Non-responders (CGI Global Improvement < "much improved")

RR (fixed) RR (fixed) Study Paroxetine Placebo or sub-category 95% CI 95% CI π/N n/N Carpenter 1999 42/96 55/98 0.78 [0.59, 1.04] Total (95% CI) 98 0.78 [0.59, 1.04] 96 Total events: 42 (Paroxetine), 55 (Placebo) Test for heterogeneity; not applicable Test for overall effect: Z = 1.70 (P = 0.09)

0.5

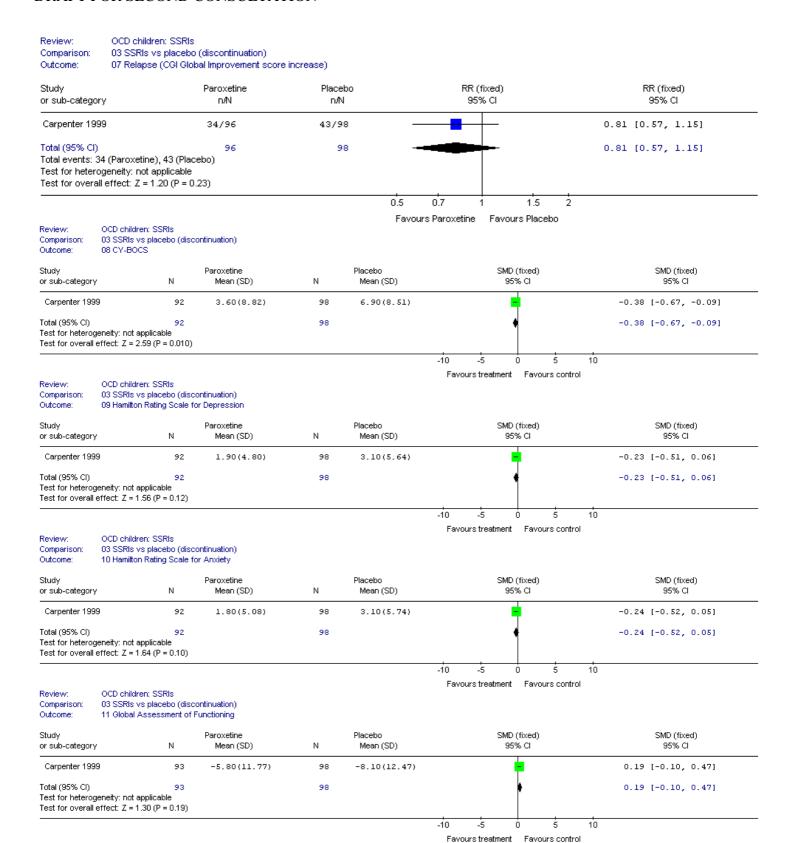
0.7

Favours Paroxetine

1.5

Favours Placebo

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Clomipramine (OCD, children)

Review: OCD children: Clomipramine Comparison: 01 Clomipramine vs placebo Outcome: 01 Leaving study early

Study or sub-category	Clomipramine n/N	Placebo n/N		RR (fixed) 95% Cl	RR (fixed) 95% Cl
DeVeaugh-Geiss 1992	4/31	2/29			1.87 [0.37, 9.46]
March 1990	2/8	0/8		-	5.00 [0.28, 90.18]
Total (95% CI)	39	37			2.48 [0.62, 9.93]
Total events: 6 (Clomipramine),	2 (Placebo)			-	
Test for heterogeneity: Chi² = 0	.34, $df = 1 (P = 0.56), I^2 = 0\%$				
Test for overall effect: Z = 1.28	(P = 0.20)				
		(0.01 0.1	1 10	100

Favours Clomipramine Favours Placebo

Favours Placebo

Favours Clomipramine Favours Placebo

Favours Clomipramine Favours Placebo

OCD children: Clomipramine 01 Clomipramine vs placebo

Comparison: 02 Leaving study early due to adverse effects Outcome:

RR (fixed) RR (fixed) Study Clomipramine Placebo or sub-category $n /\!\! N$ n/Ν 95% CI 95% CI DeVeaugh-Geiss 1992 2/31 0/29 4.69 [0.23, 93.70] March 1990 3.00 [0.14, 64.26] 1/8 0/8 Total (95% CI) 39 37 3.86 [0.45, 32.80] Total events: 3 (Clomipramine), 0 (Placebo) Test for heterogeneity: $Chi^2 = 0.04$, df = 1 (P = 0.84), $I^2 = 0\%$ Test for overall effect: Z = 1.24 (P = 0.22) 0.01 0.1 100

Favours Clomipramine

OCD children: Clomipramine Review: Comparison: 01 Clomipramine vs placebo

03 Y-BOCS

Study or sub-category	N	Clomipramine Mean (SD)	N	Placebo Mean (SD)		SMD (fix 95% (SMD (fixed) 95% Cl
March 1990	8	19.30(8.60)	8	25.60(2.40)		-		-0.94 [-1.99, 0.11]
Total (95% CI) Test for heterogeneity: not a Test for overall effect: Z = 1			8		,			-0.94 [-1.99, 0.11]
					-4	-2 0	- ;	1

OCD children: Clomipramine 01 Clomipramine vs placebo 04 NIMH-OC

Comparison: Outcome:

Study or sub-category	N	Clomipramine Mean (SD)	N	Placebo Mean (SD)			(fixed) % Cl		SMD (fixed) 95% CI
March 1990	8	7.20(2.70)	8	9.50(1.90)		-	+		-0.93 [-1.98, 0.12]
Total (95% Cl) Test for heterogeneity: not Test for overall effect: Z =			8			-	-		-0.93 [-1.98, 0.12]
					-4	-2	Ö	2	4

Review: OCD children: Clomipramine

02 Clomipramine continuation vs Desipramine substitution Comparison:

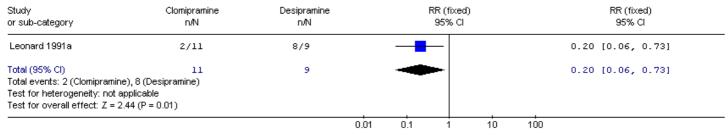
Outcome: 01 Leaving study early

Study or sub-category	Clomipramine n/N	Desipramine n/N	RR (fixed) 95% Cl	RR (fixed) 95% Cl
Leonard 1991a	0/11	1/10		0.31 [0.01, 6.74]
Total (95% CI) Total events: 0 (Clomipramir Test for heterogeneity: not Test for overall effect: Z = I	applicable	10		0.31 [0.01, 6.74]
			0.01 0.1 1 10	100
			Favours Clomipramine Favours Desiprar	nine

Review: OCD children: Clomipramine

Comparison: 02 Clomipramine continuation vs Desipramine substitution

Outcome: 02 Relapse (Physician's Relapse Scale)



Favours Clomipramine Favours Desipramine

Review: OCD children: Clomipramine

Comparison: 02 Clomipramine continuation vs Desipramine substitution

Outcome: 03 Comprehensive Psychopathological Rating Scale: obsessive-compulsive

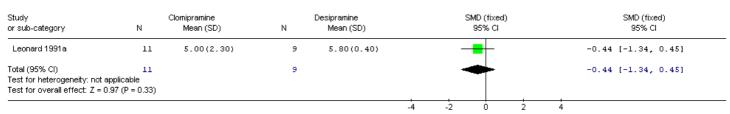
Study or sub-category	N	Clomipramine Mean (SD)	N	Desipramine Mean (SD)		SMD (fixed) 95% Cl		SMD (fixed) 95% CI
Leonard 1991a	11	5.20(4.00)	9	6.30(3.30)		-		-0.28 [-1.17, 0.60]
Total (95% CI) Test for heterogeneity: not Test for overall effect: Z =			9					-0.28 [-1.17, 0.60]
					4	· ·	<u> </u>	1

Favours Clomipramine Favours Desipramine

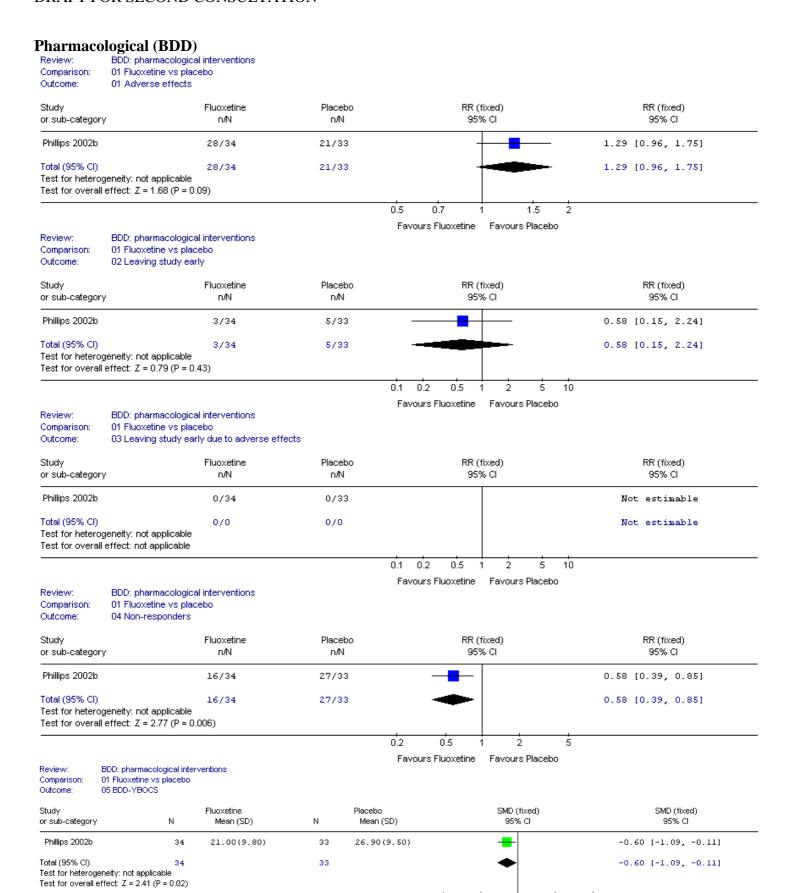
Review: OCD children: Clomipramine

Comparison: 02 Clomipramine continuation vs Desipramine substitution

Outcome: 04 NIMH-OC

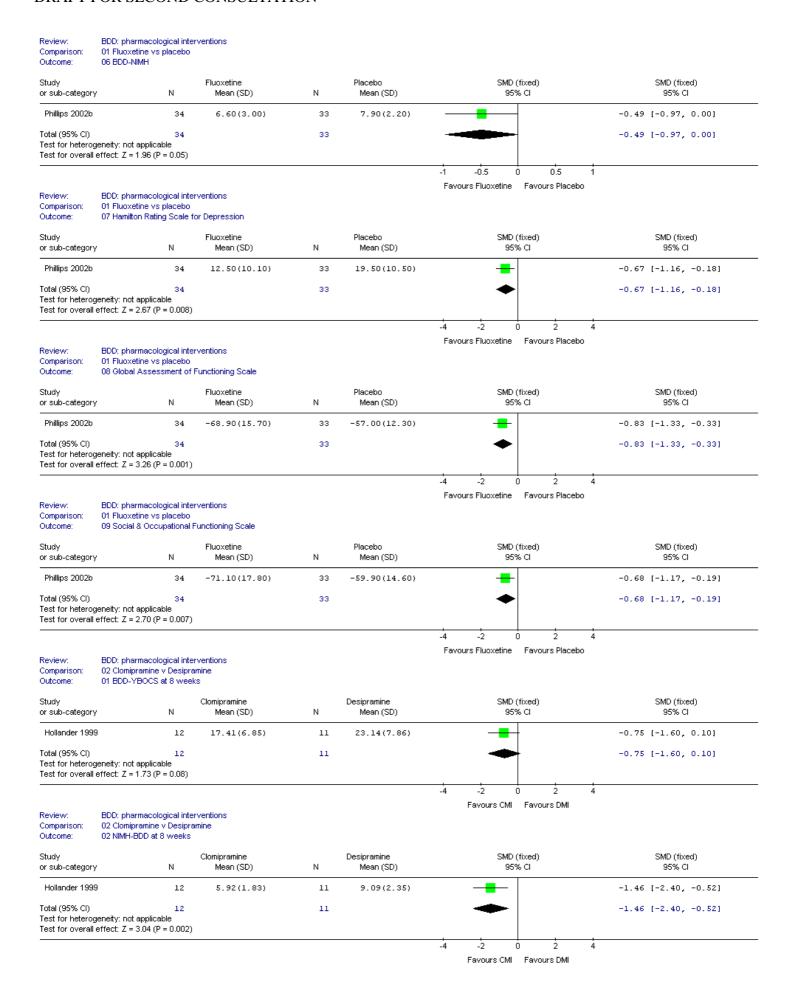


Favours Clomipramine Favours Desipramine



Favours Fluoxetine

Favours Placebo



BDD: pharmacological interventions Review: Comparison: 02 Clomipramine v Desipramine 03 HAM-D at 8 weeks Outcome:

Desipramine SMD (fixed) SMD (fixed) Study Clomipramine or sub-category N Mean (SD) Ν Mean (SD) 95% CI 95% CI Hollander 1999 9.42(7.89) 12 11 14.64(5.61) -0.73 [-1.58, 0.12] Total (95% CI) 11 -0.73 [-1.58, 0.12] Test for heterogeneity: not applicable Test for overall effect: Z = 1.68 (P = 0.09) Ó

Favours CMI Favours DMI

Favours Placebo

Favours Placebo

Favours Pimozide Favours Placebo

Review: BDD: pharmacological interventions

Comparison: 03 Pimozide v Placebo

Outcome: 01 Non-responders (Y-BOCS 30%)

Study or sub-category	Pimozide n/N	Placebo n/N		(fixed) 5% Cl		RR (fixed) 95% Cl
Phillips_in press	9/11	14/17	_	+	(0.99 [0.70, 1.42]
Total (95% CI) Test for heterogeneity: not ap Test for overall effect: Z = 0.0	•	14/17		+		0.99 [0.70, 1.42]
		0.2	0.5	1 2	5	

Favours Pimozide

Favours Pimozide

Review: BDD: pharmacological interventions

03 Pimozide v Placebo Comparison: Outcome: 02 Leaving the study early

RR (fixed) RR (fixed) Study Pimozide Placebo 95% CI or sub-category $n /\!\! N$ n/N 95% CI Phillips_in press 6/11 13/17 0.71 [0.39, 1.30] Total (95% CI) 6/11 13/17 0.71 [0.39, 1.30] Test for heterogeneity: not applicable Test for overall effect: Z = 1.10 (P = 0.27)0.2 0.5 5

Review: BDD: pharmacological interventions

Comparison: 03 Pimozide v Placebo

Outcome 03 Y-BOCS

Study or sub-category	N	Pimozide Mean (SD)	N	Placebo Mean (SD)			1D (random 95% CI)		VVMD (random) 95% CI
Phillips_in press	11	23.90(6.60)	17	28.00(7.80)		-	-			-4.10 [-9.48, 1.28]
Total (95% CI) Test for heterogeneity: not Test for overall effect: Z =			17							-4.10 [-9.48, 1.28]
					-10	-5	-	5	10	

BDD: pharmacological interventions 03 Pimozide v Placebo Review:

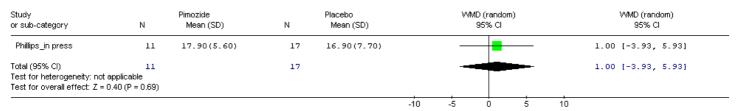
Comparison:

Outcome 04 Brown Assessment of Beliefs Scale

Study or sub-category	N	Pirnozide Mean (SD)	N	Placebo Mean (SD)	VVMD (random) 95% Cl	VMMD (random) 95% CI
Phillips_in press	11	13.50(7.30)	17	17.20(6.00)		-3.70 [-8.87, 1.47]
Total (95% CI) Test for heterogeneity: not Test for overall effect: Z =	• •		17			-3.70 [-8.87, 1.47]
					-10 -5 0 5 Favours Pimozide Favours Placeh	10

Review: BDD: pharmacological interventions Comparison: 03 Pimozide v Placebo

Outcome: 05 Hamilton Depression Rating Scale



Favours Pimozide Favours Placebo

Review: BDD: pharmacological interventions

Comparison: 03 Pimozide v Placebo Outcome: 06 Brief Psychiatric Rating Scale

