

## Appendix 19: Clinical evidence forest plots

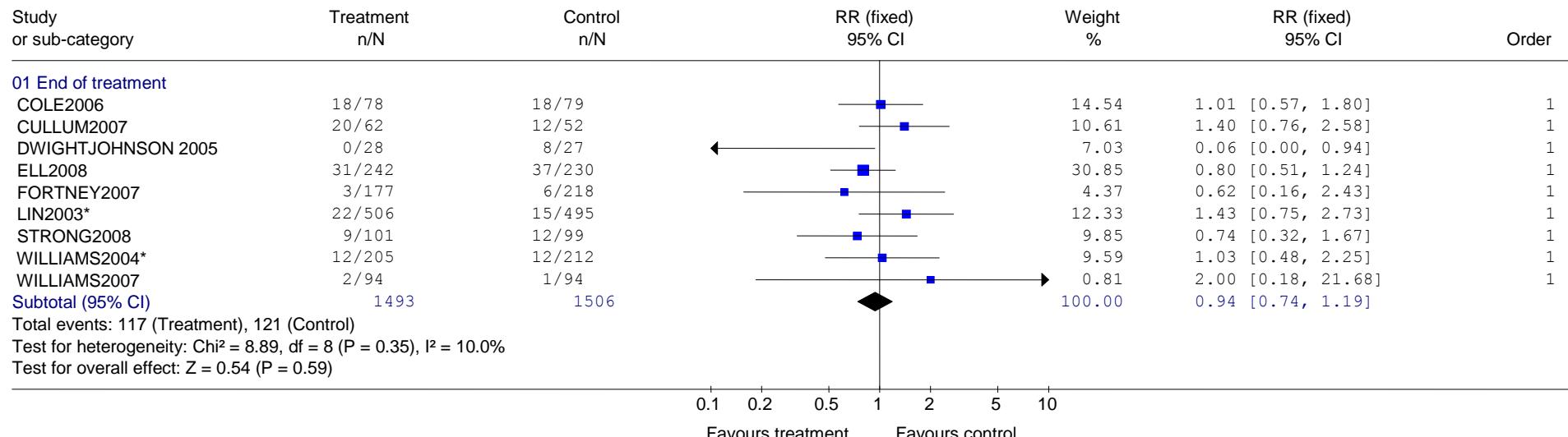
Service-level interventions.....	3
Collaborative care versus everything .....	3
Collaborative care - subgroup analysis by comparison (standard care, enhanced standard care).....	16
Collaborative care - sensitivity analysis: ICC calculations.....	25
Collaborative care - sensitivity analysis: recruited for comorbid physical health condition.....	34
Other service-level interventions versus standard care .....	41
Psychological and psychosocial interventions .....	44
Physical activity versus standard care .....	44
Peer (self-help) support versus standard care .....	48
Peer (self-help) support versus group based cognitive and behavioural interventions .....	50
Guided self-help based on cognitive and behavioural principles versus standard care .....	51
Individual guided self-help based on McMaster model of family functioning versus standard care .....	52
Health education versus standard care .....	53
Relaxation training versus standard care .....	54
Social support versus standard care .....	56
Individual-based cognitive and behavioural interventions versus standard care.....	56
Individual-based cognitive and behavioural interventions versus counselling .....	59
Group-based cognitive and behavioural interventions versus standard care.....	61
Group-based cognitive and behavioural interventions versus other (peer self-help support and health education) .....	66
Counselling versus standard care .....	67
Group existential therapy versus active control .....	68
Combination interventions.....	70
Psychological/psychosocial (psych) interventions plus pharmacological interventions (pharm) versus psychological/psychosocial intervention alone .....	70
Psychological/psychosocial interventions (psych) plus pharmacological interventions (pharm) versus pharmacological interventions alone.....	72
Psychological/psychosocial versus pharmacological interventions.....	73
Psychological/psychosocial interventions (psych) versus pharmacological interventions (pharm).....	73
Pharmacological interventions .....	74
SSRIs versus placebo .....	74

TCAs versus placebo .....	93
Third-generation drugs versus placebo.....	100
Mianserin versus placebo .....	103
Trazodone versus placebo .....	106
Any antidepressant versus placebo – sensitivity analysis removing non-blinded studies .....	107
Any antidepressant versus placebo – sensitivity analysis removing subthreshold studies .....	110
Any antidepressant versus placebo – subgroup analysis: HAM-D only.....	113
SSRIs versus TCAs.....	115
SSRI versus placebo – sensitivity analysis removing non-blinded and subthreshold studies.....	125
TCAs versus placebo – sensitivity analysis removing non-blinded and subthreshold studies.....	129

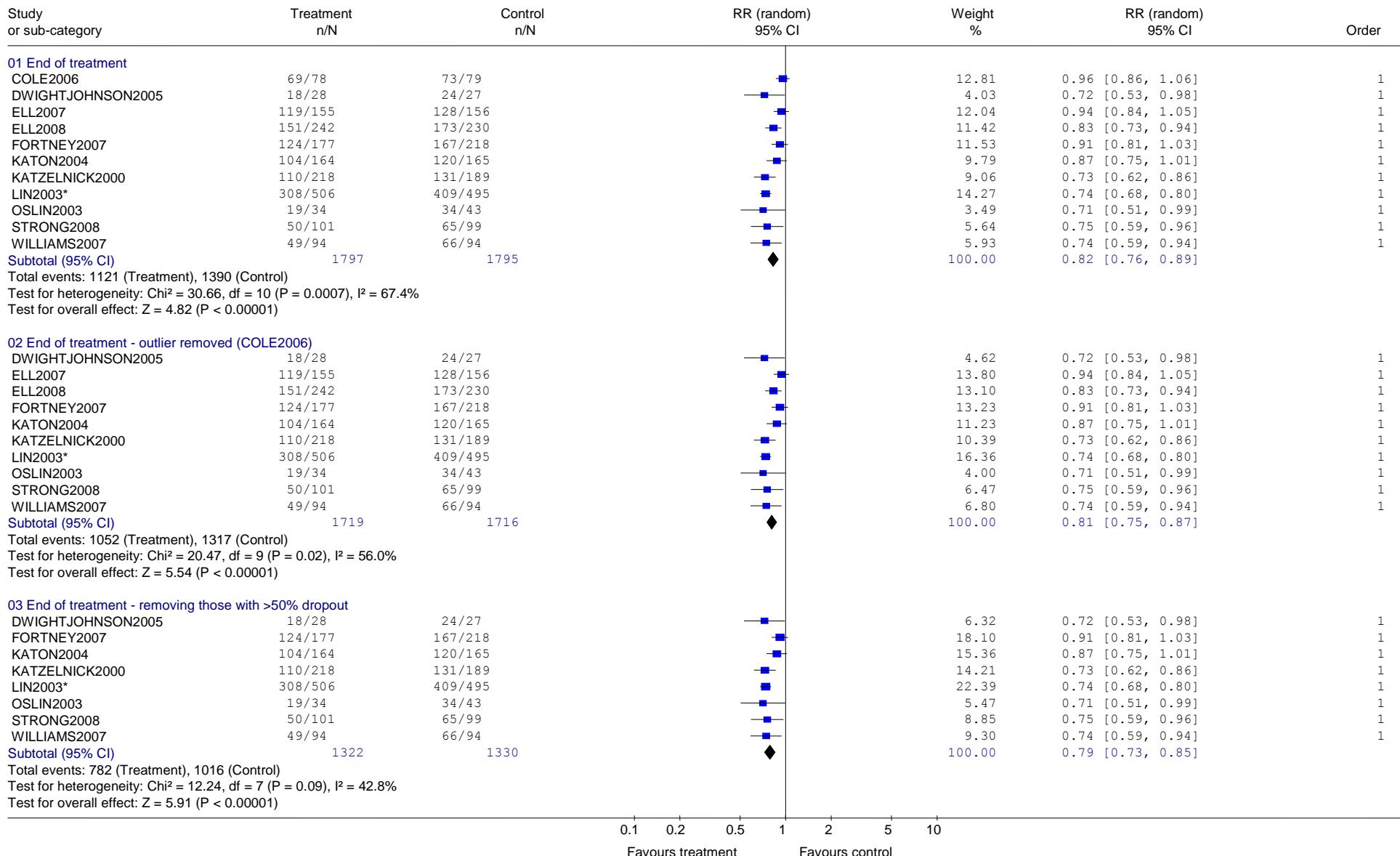
## *Service-level interventions*

### **Collaborative care versus everything**

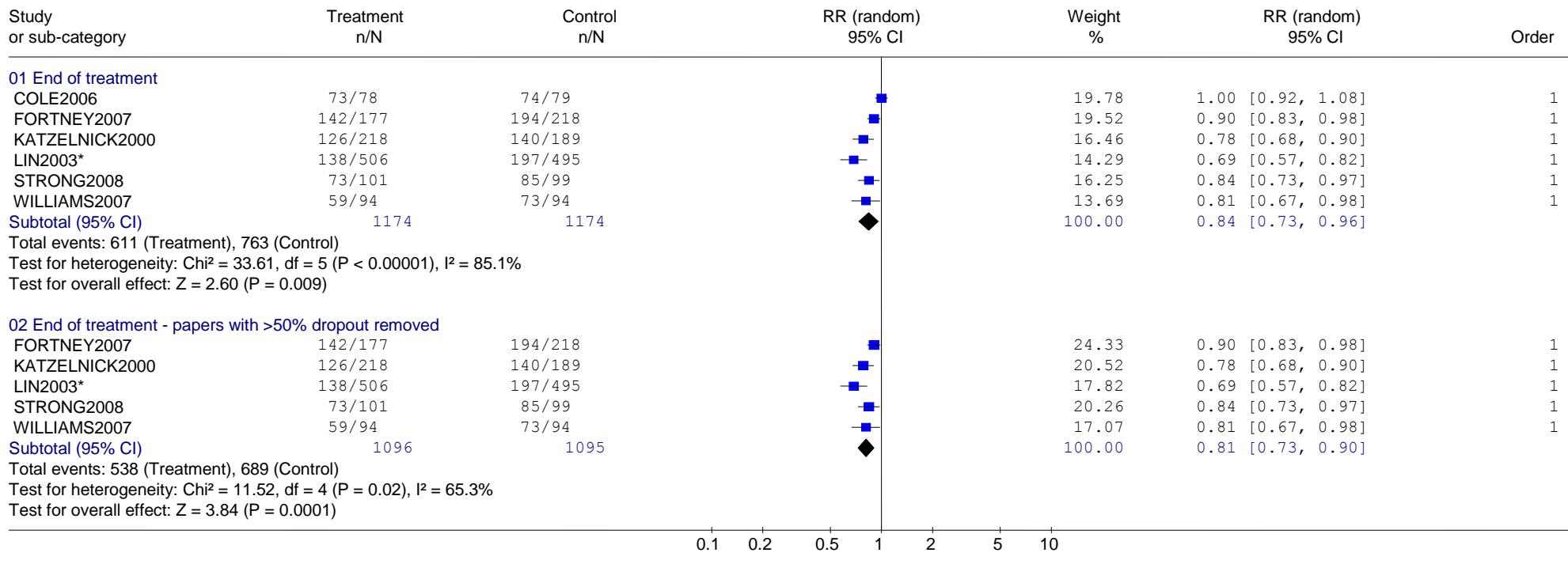
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 20 Mortality



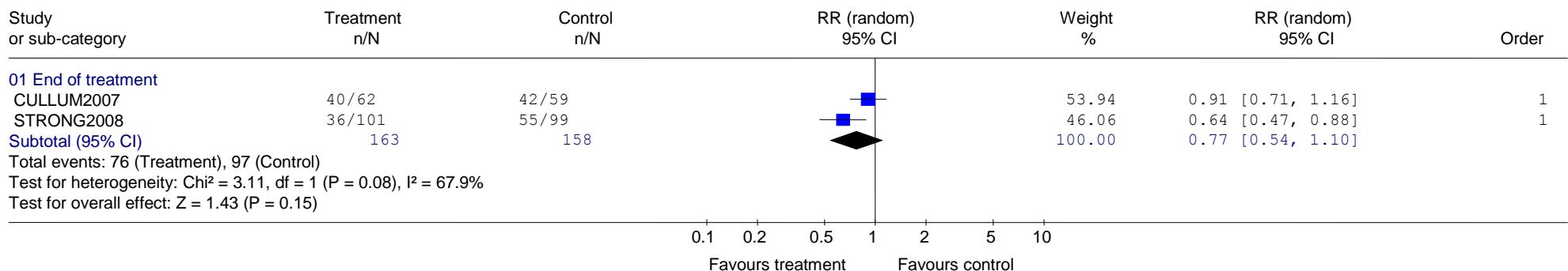
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 30 Depression outcome. 1. Non-response (<50% improvement) - sensitivity analysis



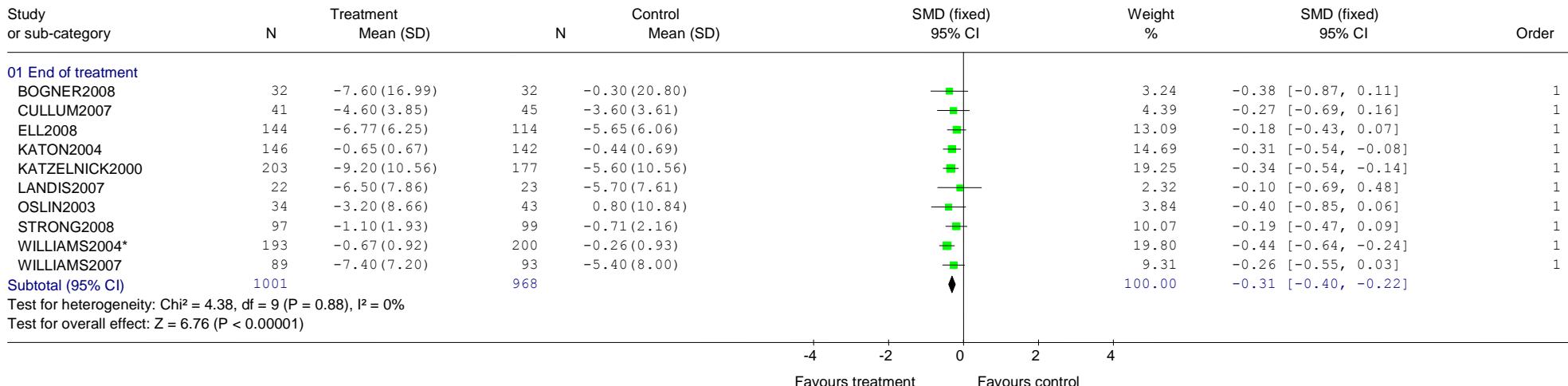
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 32 Depression outcome 2. Non-remission (scoring above cut-off, for example >7 on HAM-D)



Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 33 Depression outcome 3. Diagnosis (at follow-up)



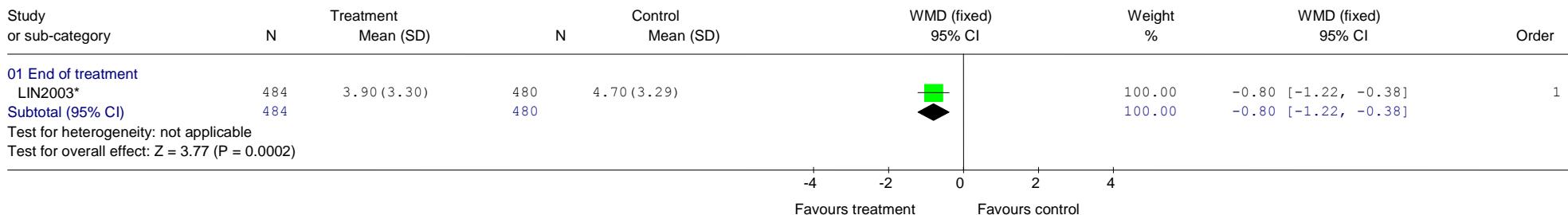
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 34 Depression outcome 4. Continuous measures depression rating scale (Change score)



Test for heterogeneity:  $\chi^2 = 4.38$ ,  $df = 9$  ( $P = 0.88$ ),  $I^2 = 0\%$

Test for overall effect:  $Z = 6.76$  ( $P < 0.00001$ )

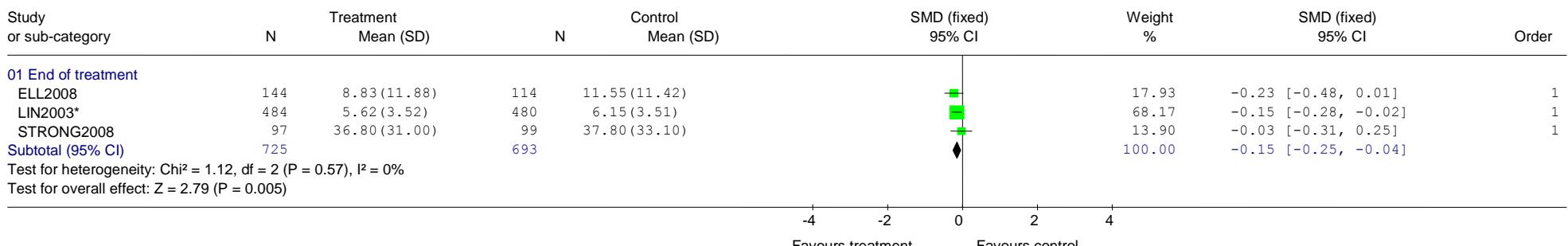
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 40 Physical health outcome: 1. Sheehan disability score



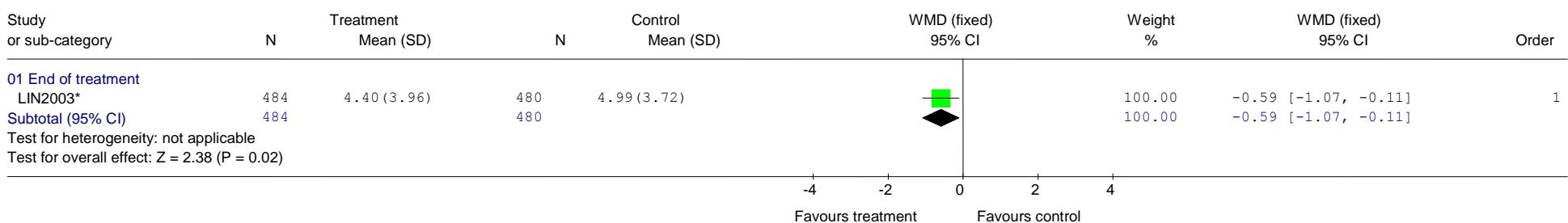
Test for heterogeneity: not applicable

Test for overall effect:  $Z = 3.77$  ( $P = 0.0002$ )

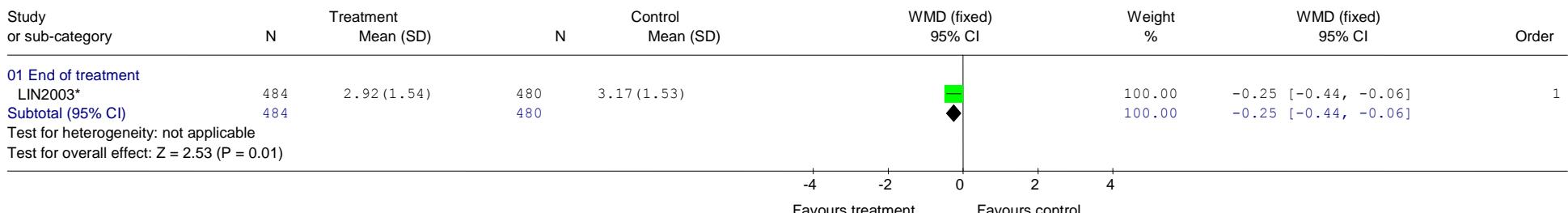
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 41 Physical health outcome: 2. Pain intensity (Brief Pain Inventory, author defined scale)



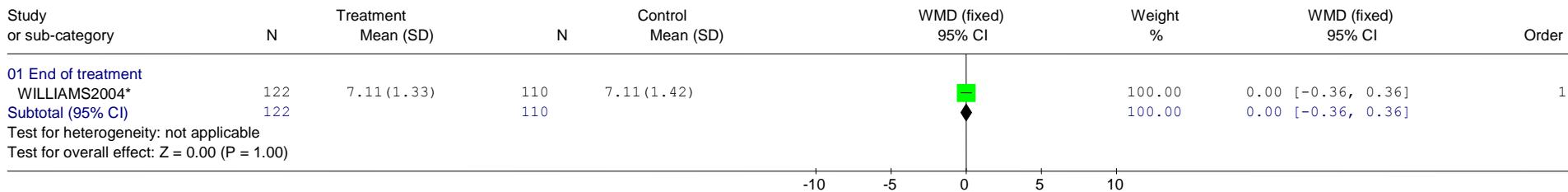
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 42 Physical health outcome: 3. Interference with daily activities



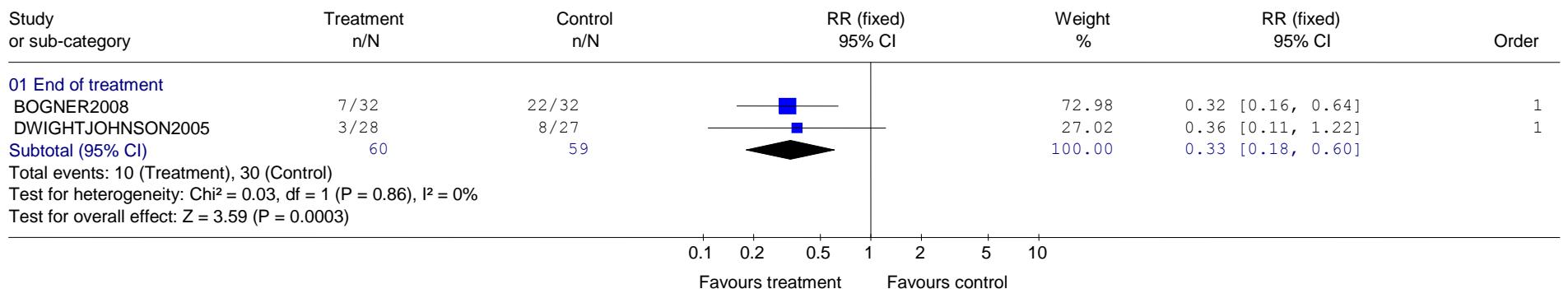
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 43 Physical health outcome: 4. Pain interferes with activity



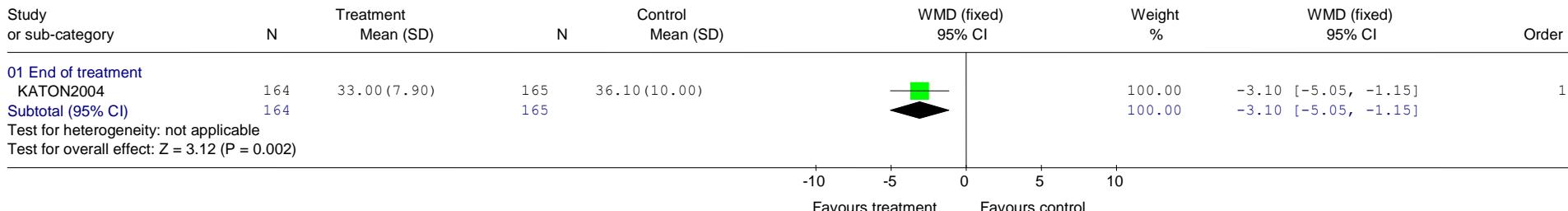
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 44 Physical health outcome: 5. Haemoglobin values



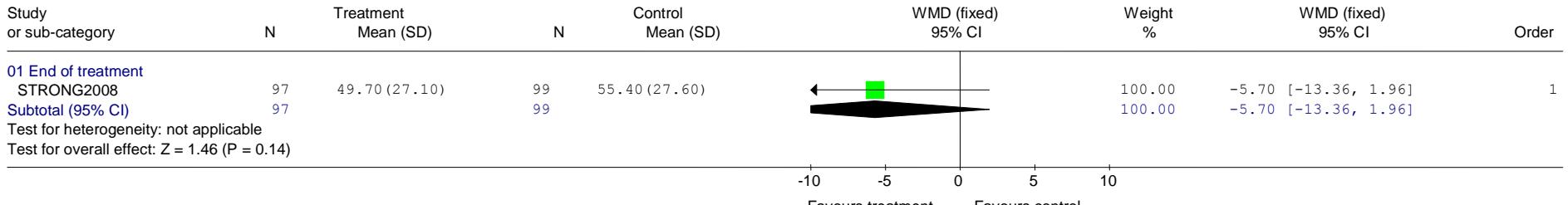
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 45 Physical health outcome 6. Non-adherence to physical health treatment



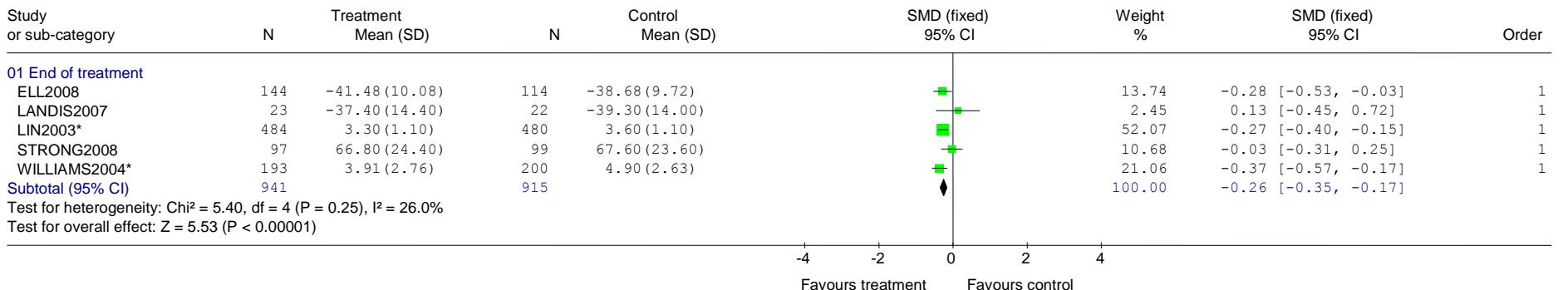
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 46 Physical health outcome: 7. BMI



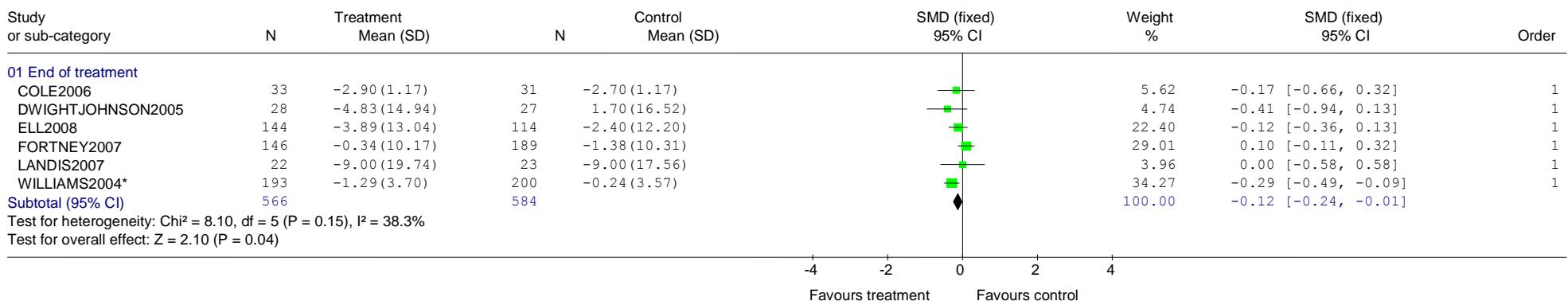
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 47 Physical health outcome: 8. Fatigue



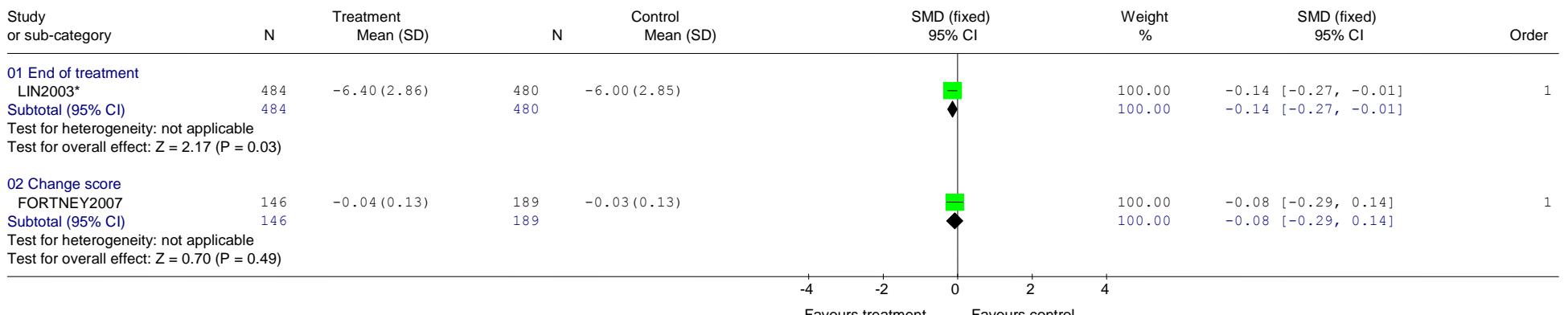
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 49 Physical health outcome/ QoL: 1. General physical wellbeing/ functioning (SF-12 physical etc)



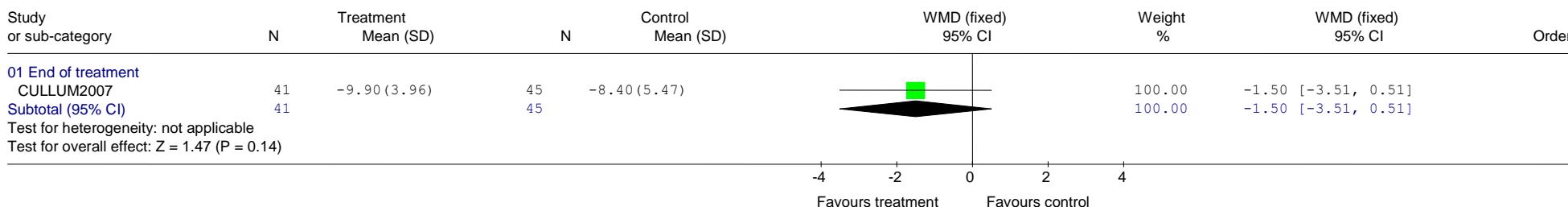
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 50 Physical health outcome/ QoL: 2. General physical wellbeing/ functioning (Change scores)



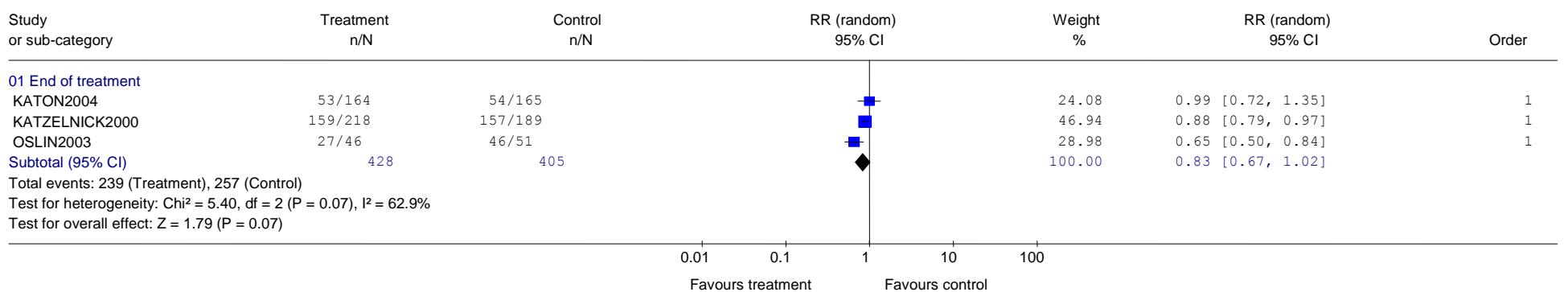
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 55 Quality of Life 1. General QoL scales (Euroqol, 0-10 rating scale etc.)



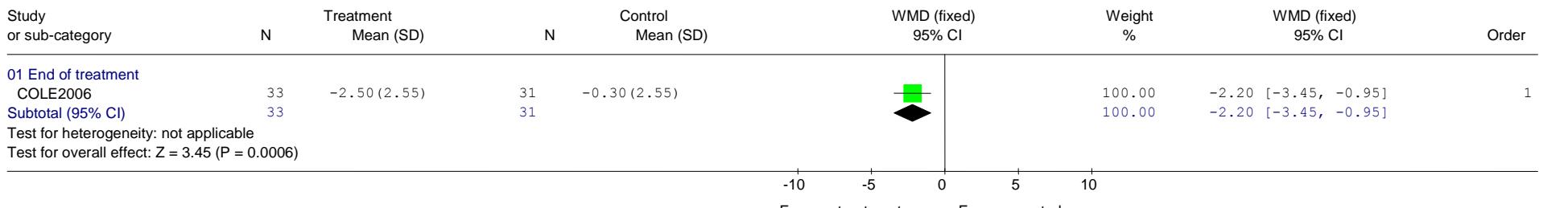
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 56 Quality of Life 2. No. QALWs gained (signs reversed)



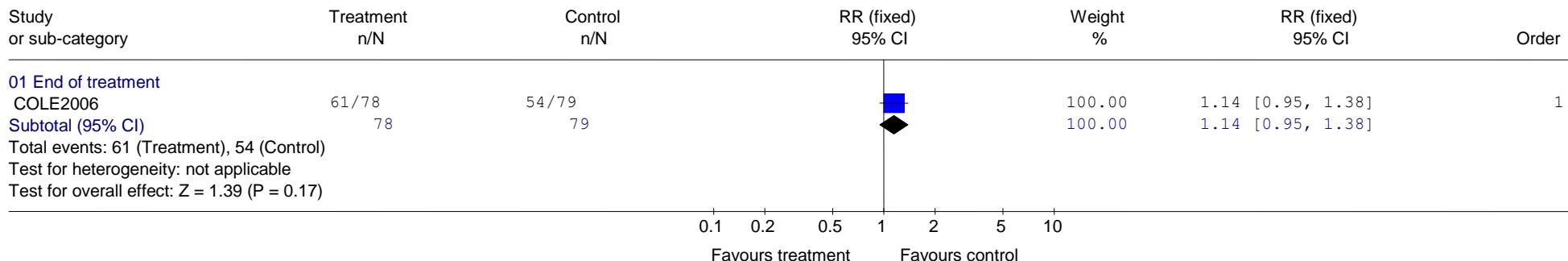
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 60 Service use/ Process of care: 1. Did not receive a consultation / specified number of mental health visits



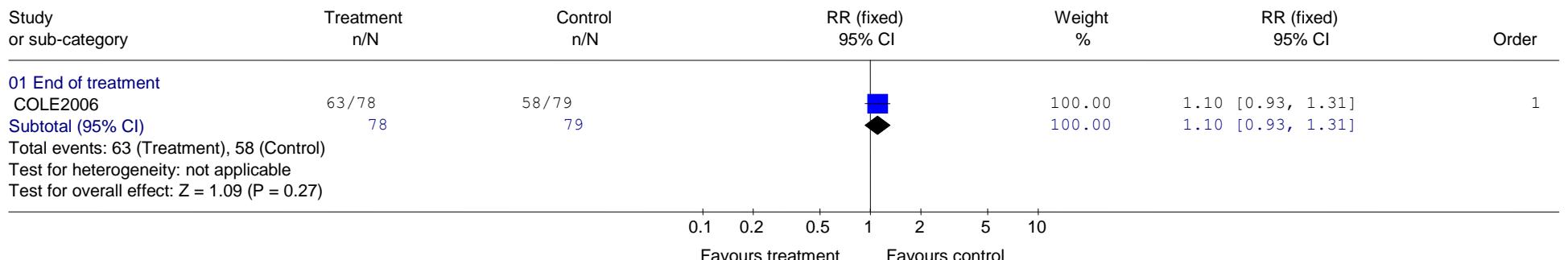
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 61 Service use/ Process of care: 2. Number of psychiatric outpatient visits



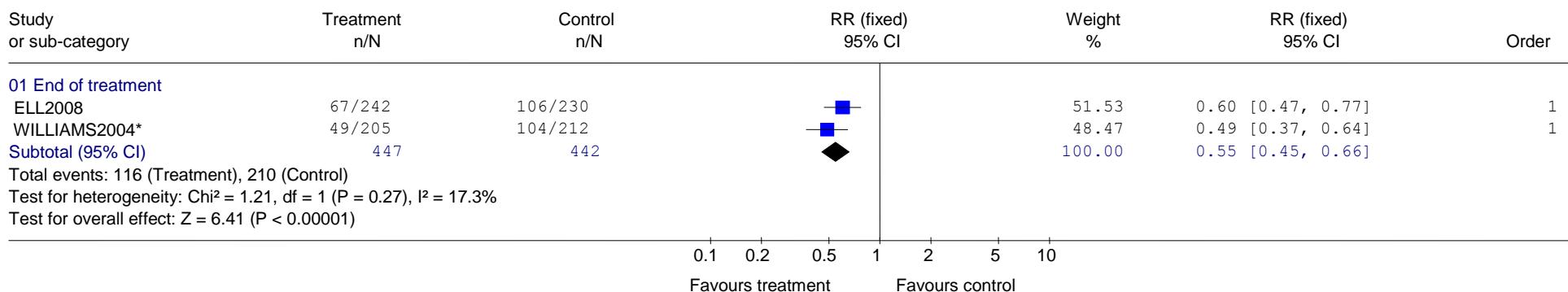
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 62 Service use/ Process of care: 3. Readmission



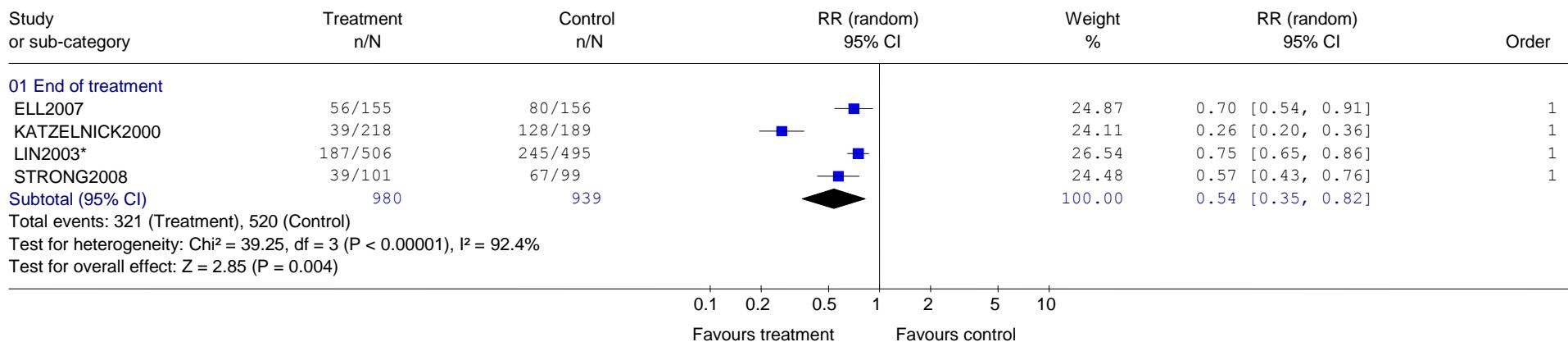
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 63 Service use/ Process of care: 4. Any visit to A&E



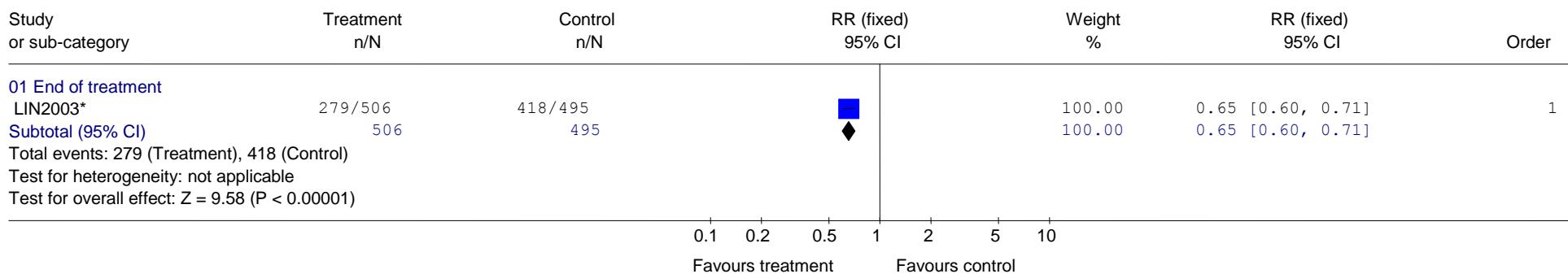
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 64 Service use/ process of care : 5. Not receiving any depression treatment (antidepressants or psychosocial)



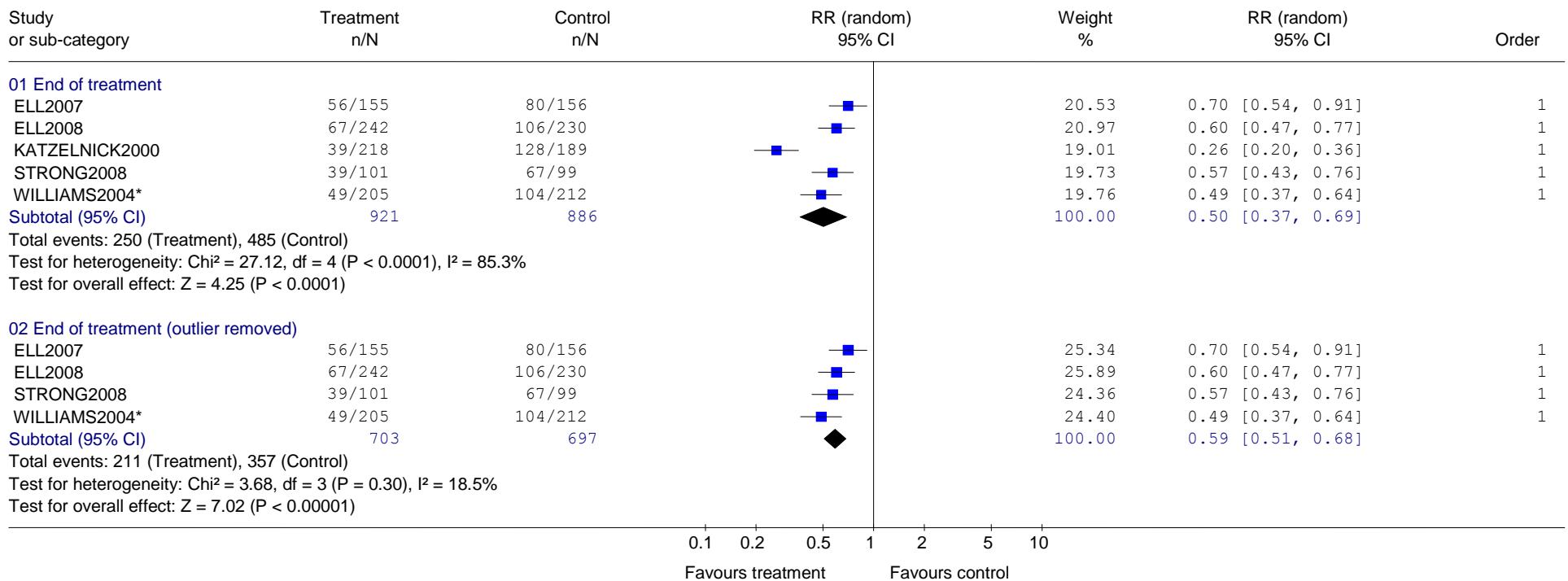
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 65 Service use/ process of care : 6. Not receiving antidepressant medication



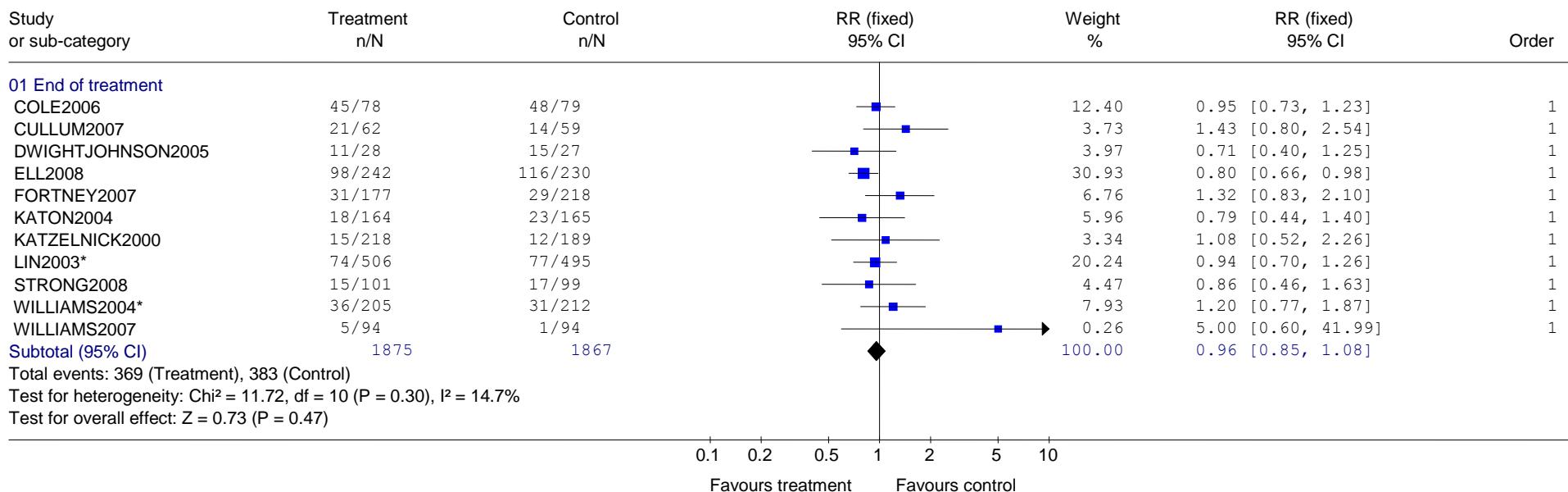
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 66 Service use/ process of care : 7. Not receiving psychosocial interventions



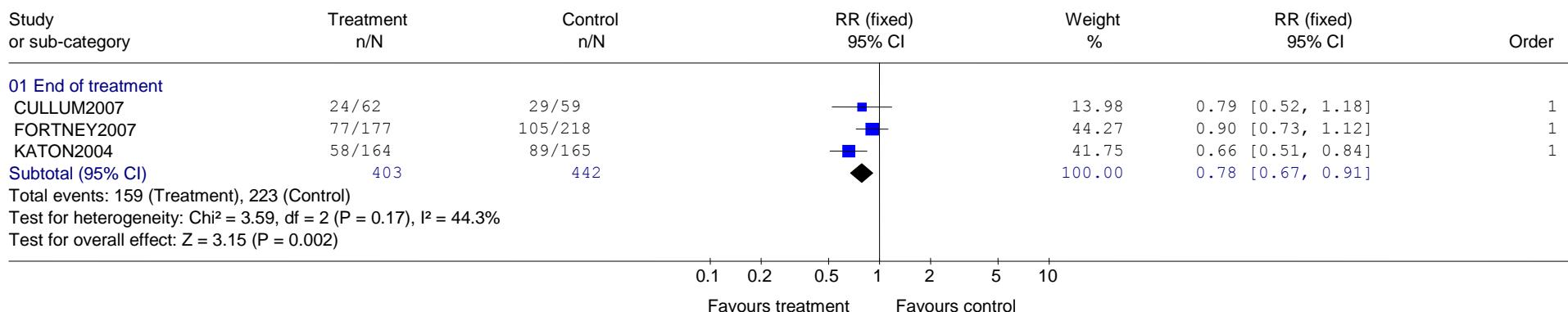
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 67 Service use / Process of care (64-66 combined)



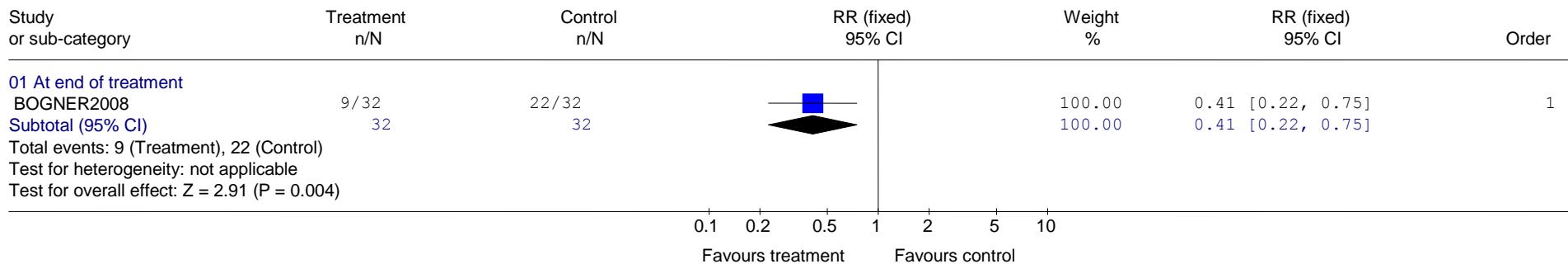
Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 70 Treatment acceptability - leaving the study early for any reason



Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 80 Satisfaction with service - not satisfied with treatment/care

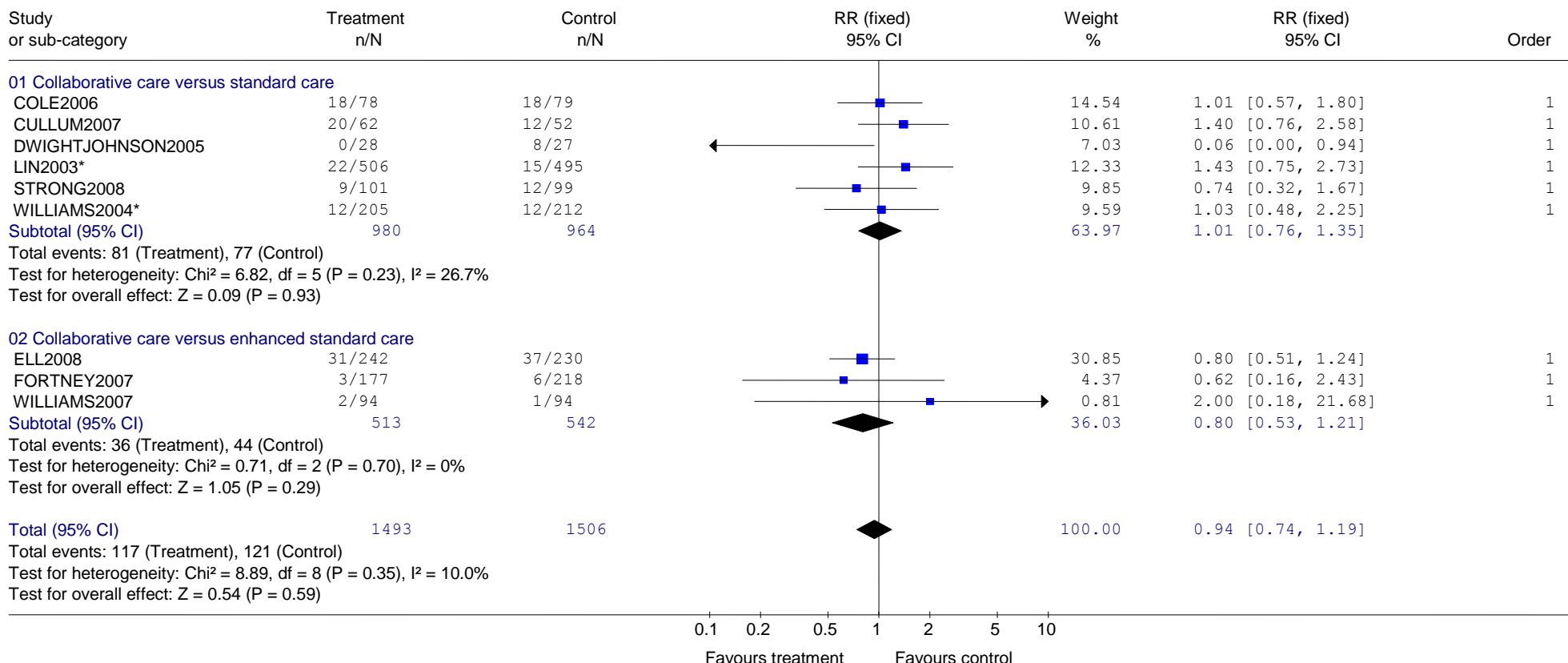


Review: DCHP - Service  
 Comparison: 01 Collaborative care versus everything (WCS analysis)  
 Outcome: 81 Non-adherence to antidepressant medication

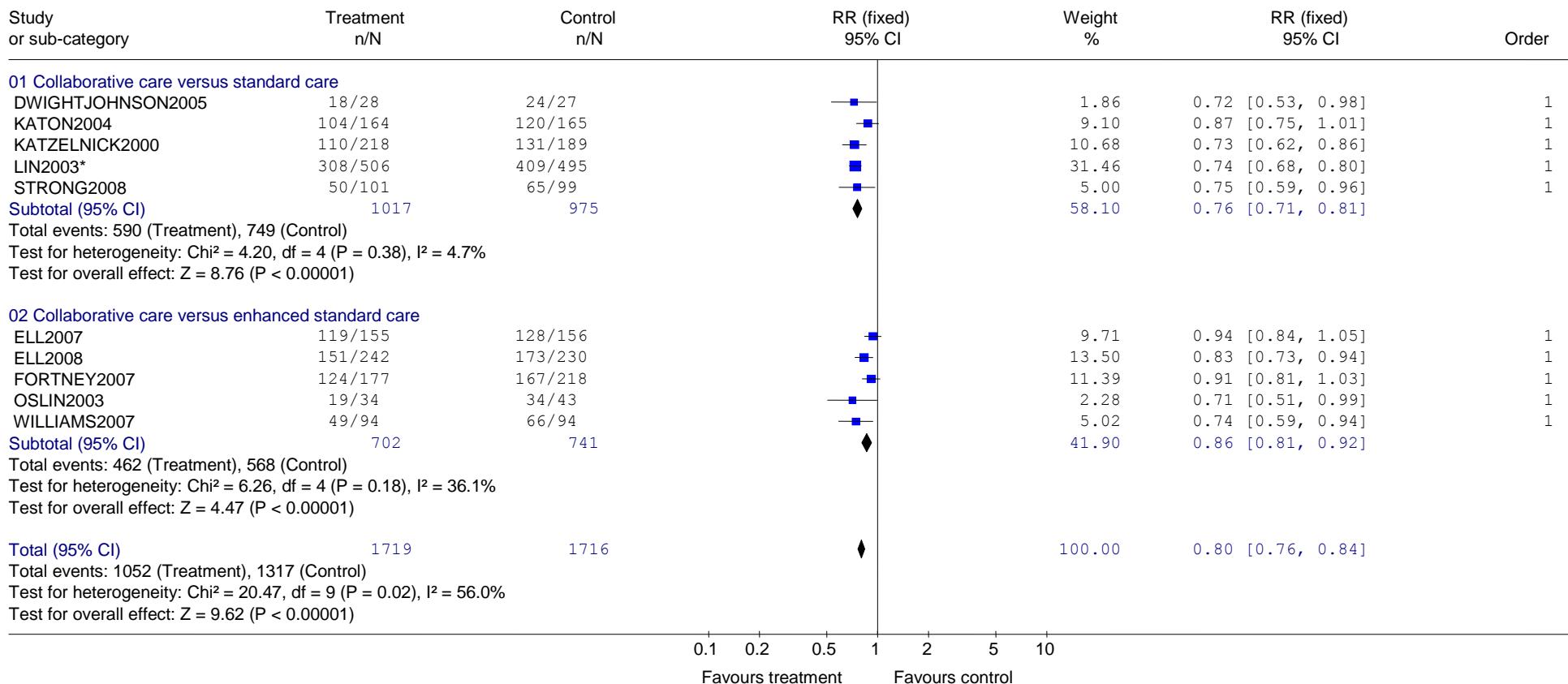


## Collaborative care – subgroup analysis by comparison (standard care, enhanced standard care)

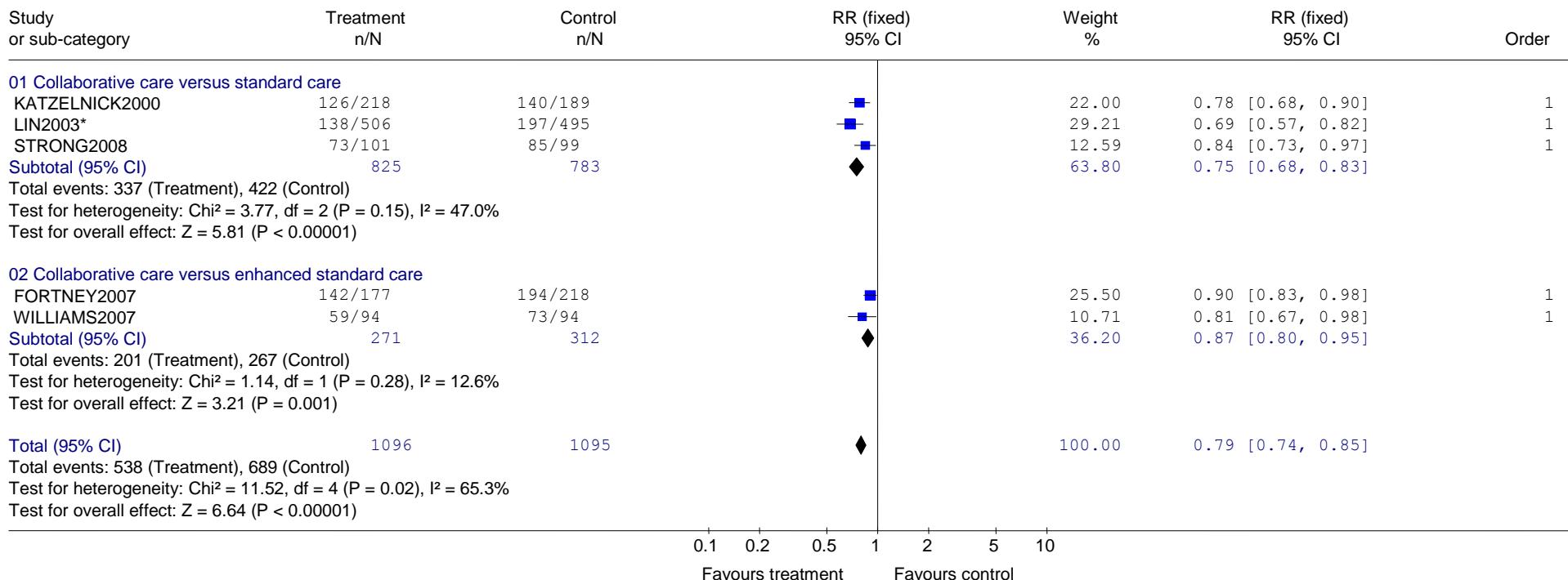
Review: DCHP - Service  
 Comparison: 02 Collaborative care vs. everything (WCS analysis) - subgroup by comparison  
 Outcome: 20 Mortality



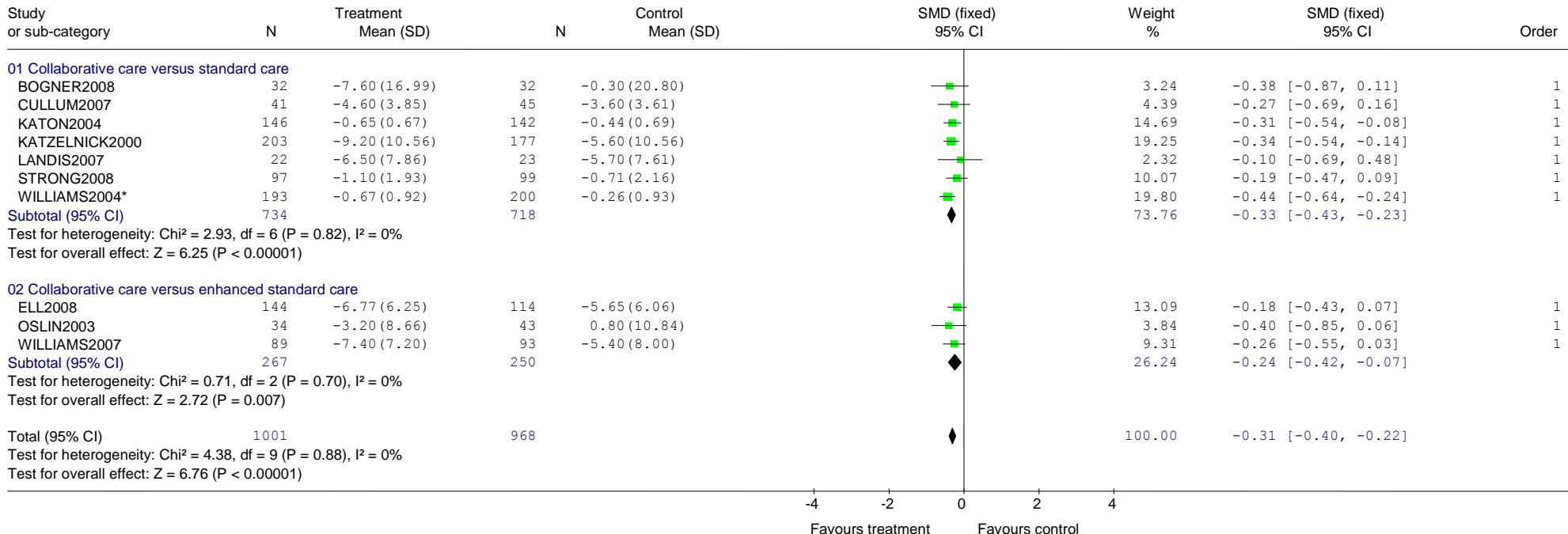
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 31 Depression outcome: 1. Non-Response (<50% improvement) - removing outlier



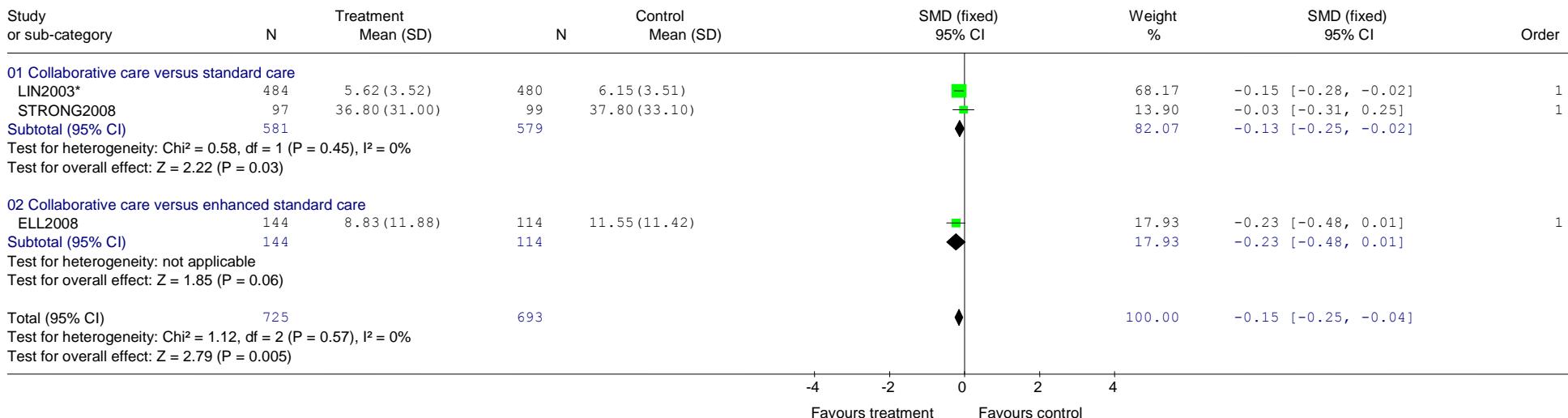
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 32 Depression outcome: 2. Non-remission (scoring above cut-off, for example >7 on HAM-D) - removed outlier



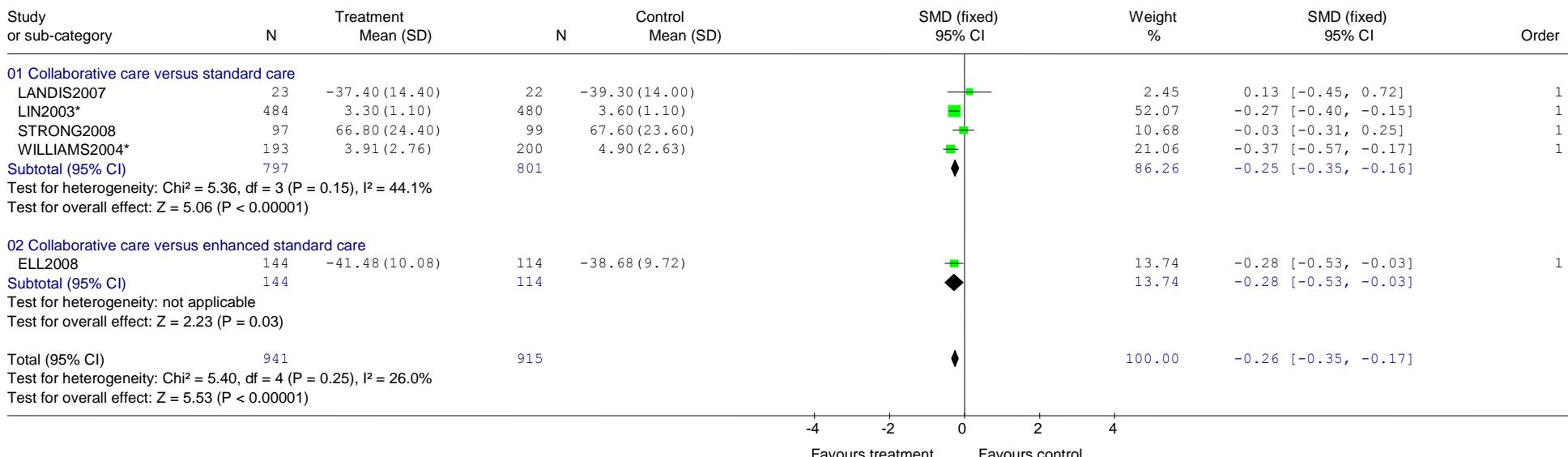
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 34 Depression outcome: 4. Continuous measures depression rating scale (Change score)



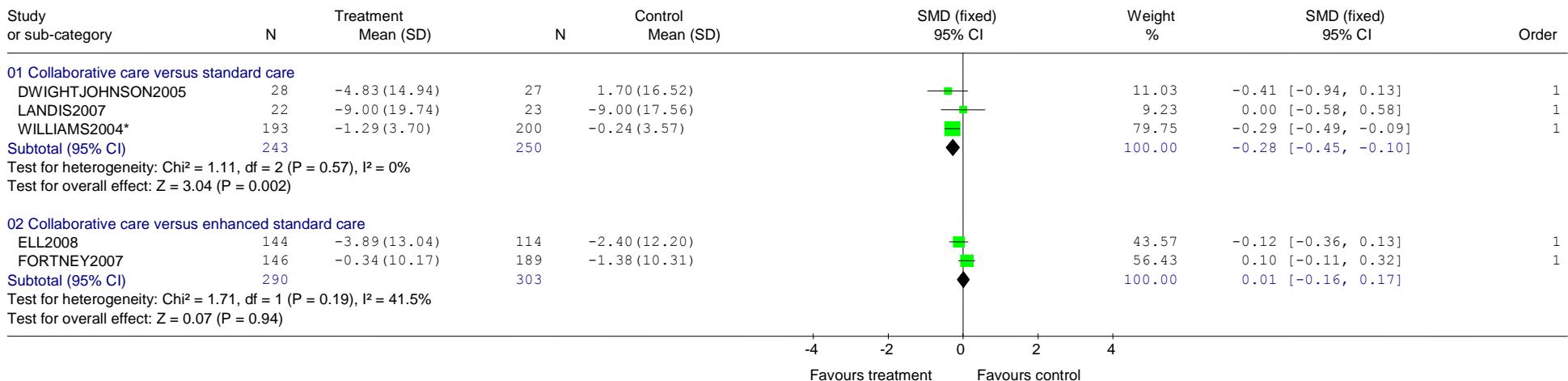
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 41 Physical health outcome: 2. Pain intensity (Brief Pain Inventory, author defined scale)



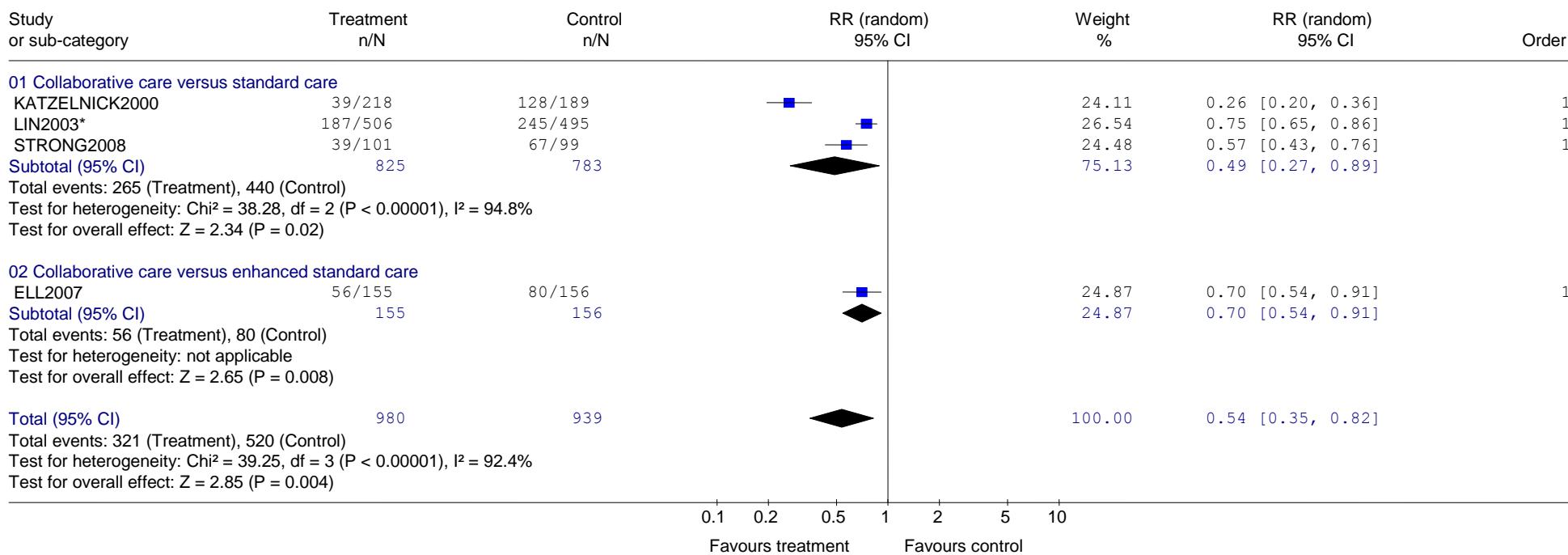
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 48 Physical health outcome/ QoL: 1. General physical well-being/ functioning (SF-12 physical etc)



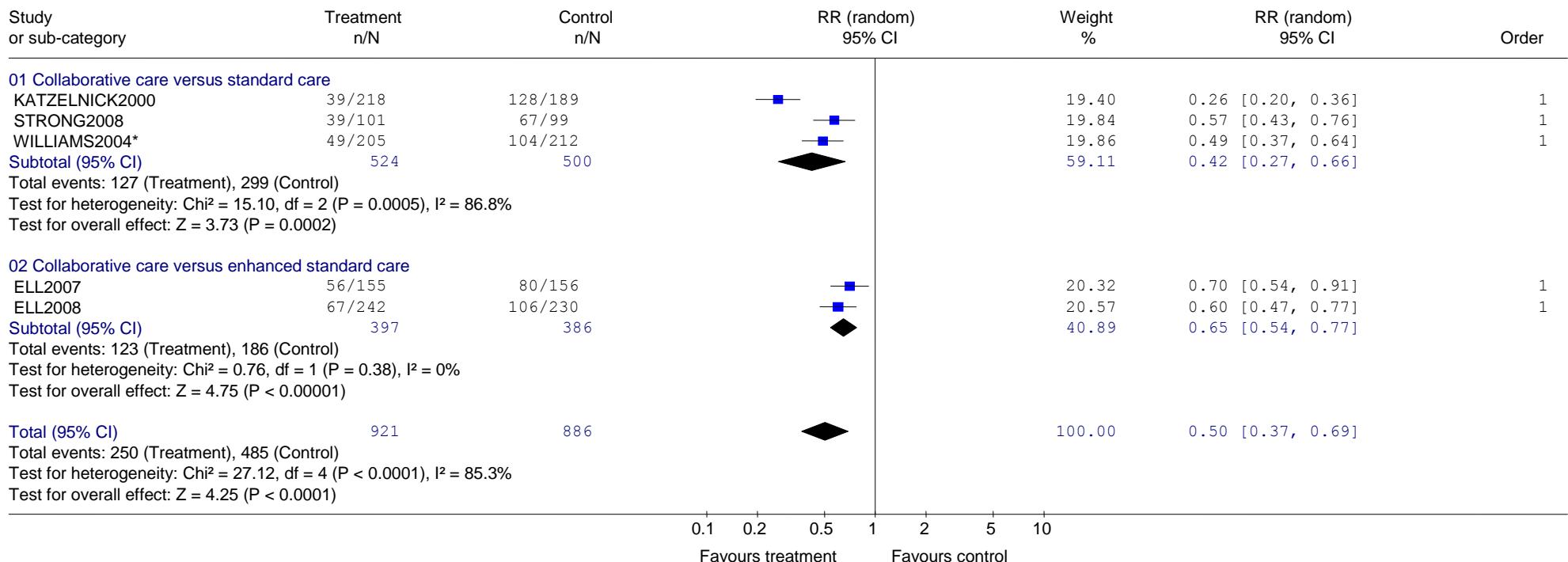
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 49 Physical health outcome/ QoL: 2. General physical well-being/ functioning (Change scores)



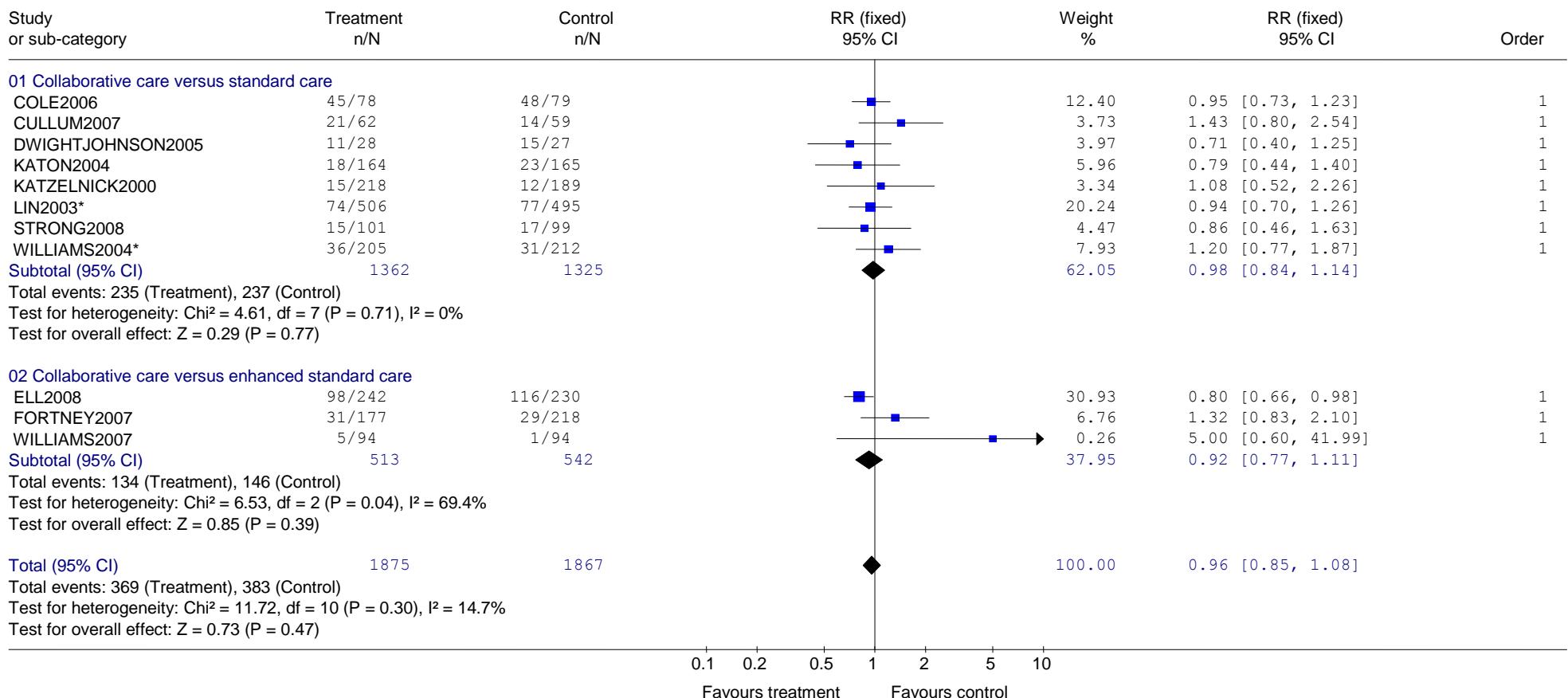
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 65 Service use/ process of care : 6. Not receiving antidepressant medication



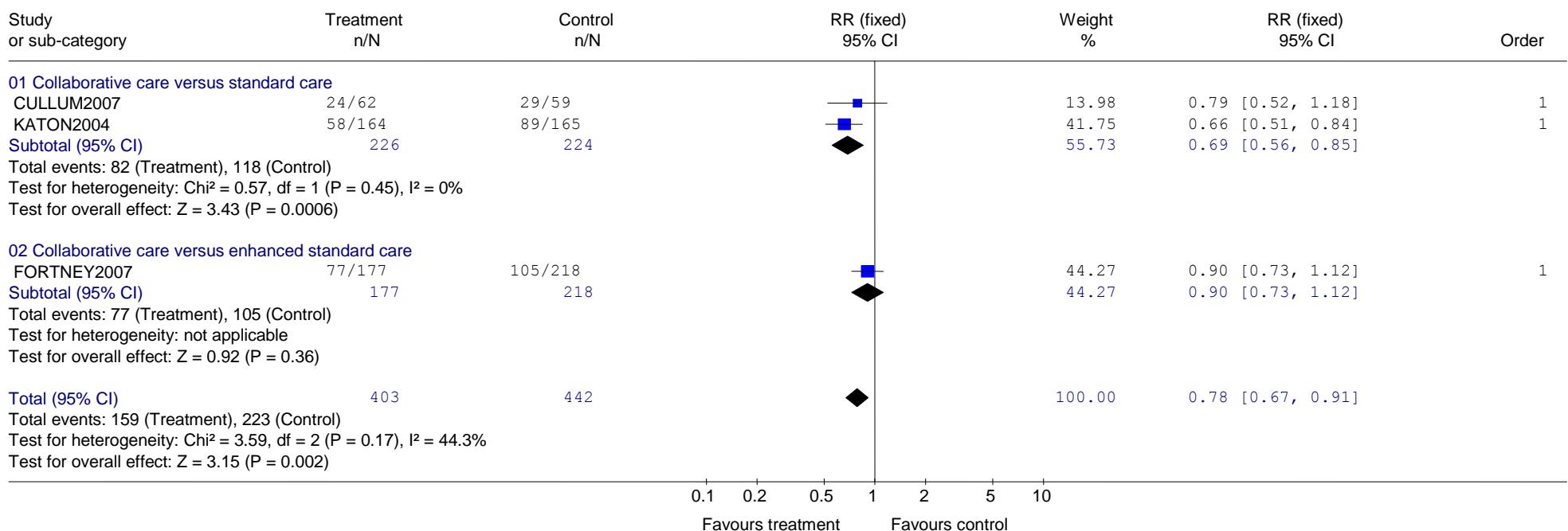
Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 67 Service use / Process of care (64-66 combined)



Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 70 Treatment acceptability - leaving the study early for any reason

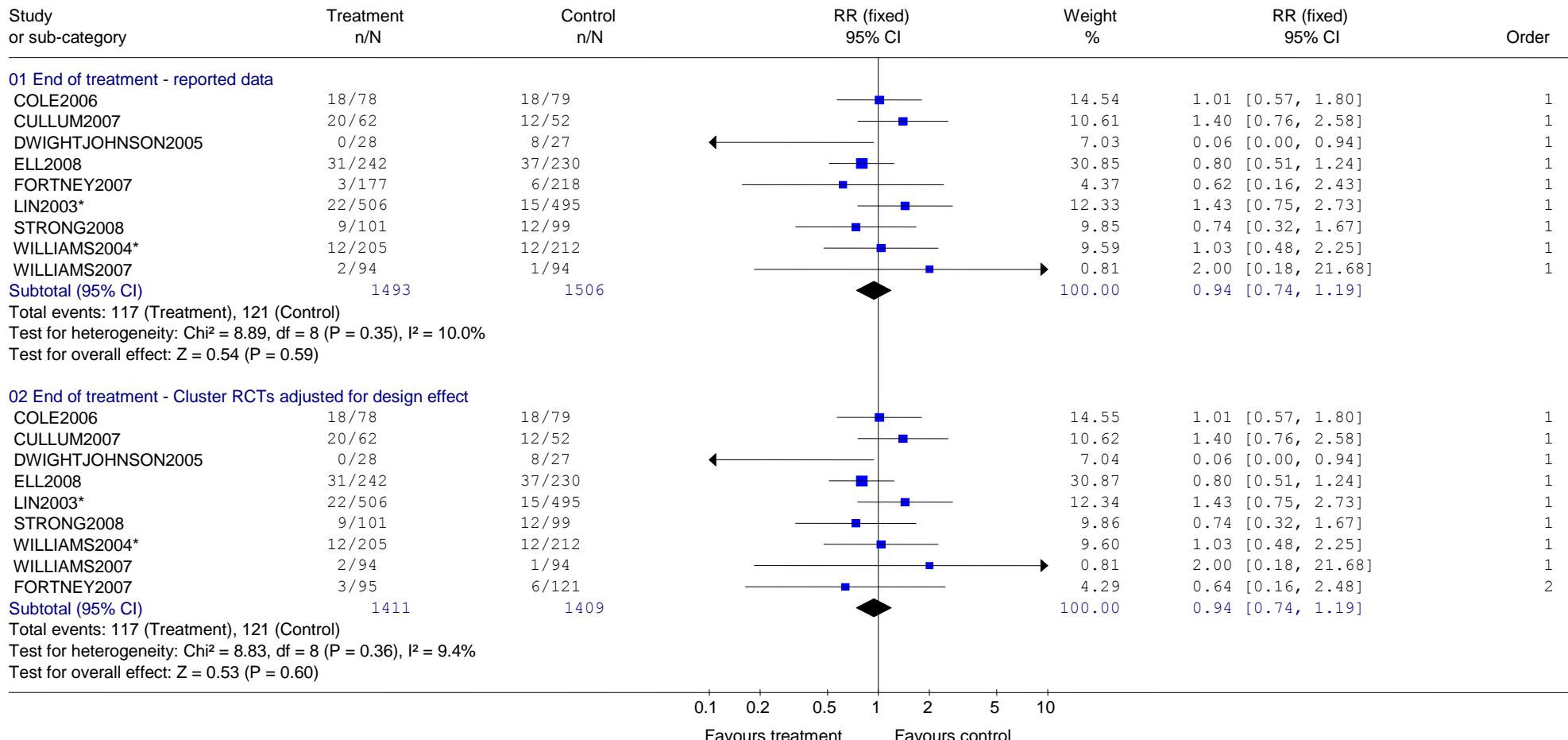


Review: DCHP - Service  
 Comparison: 02 Collaborative care versus everything (WCS analysis) - subgroup by comparison  
 Outcome: 80 Satisfaction with service - not satisfied with treatment/care

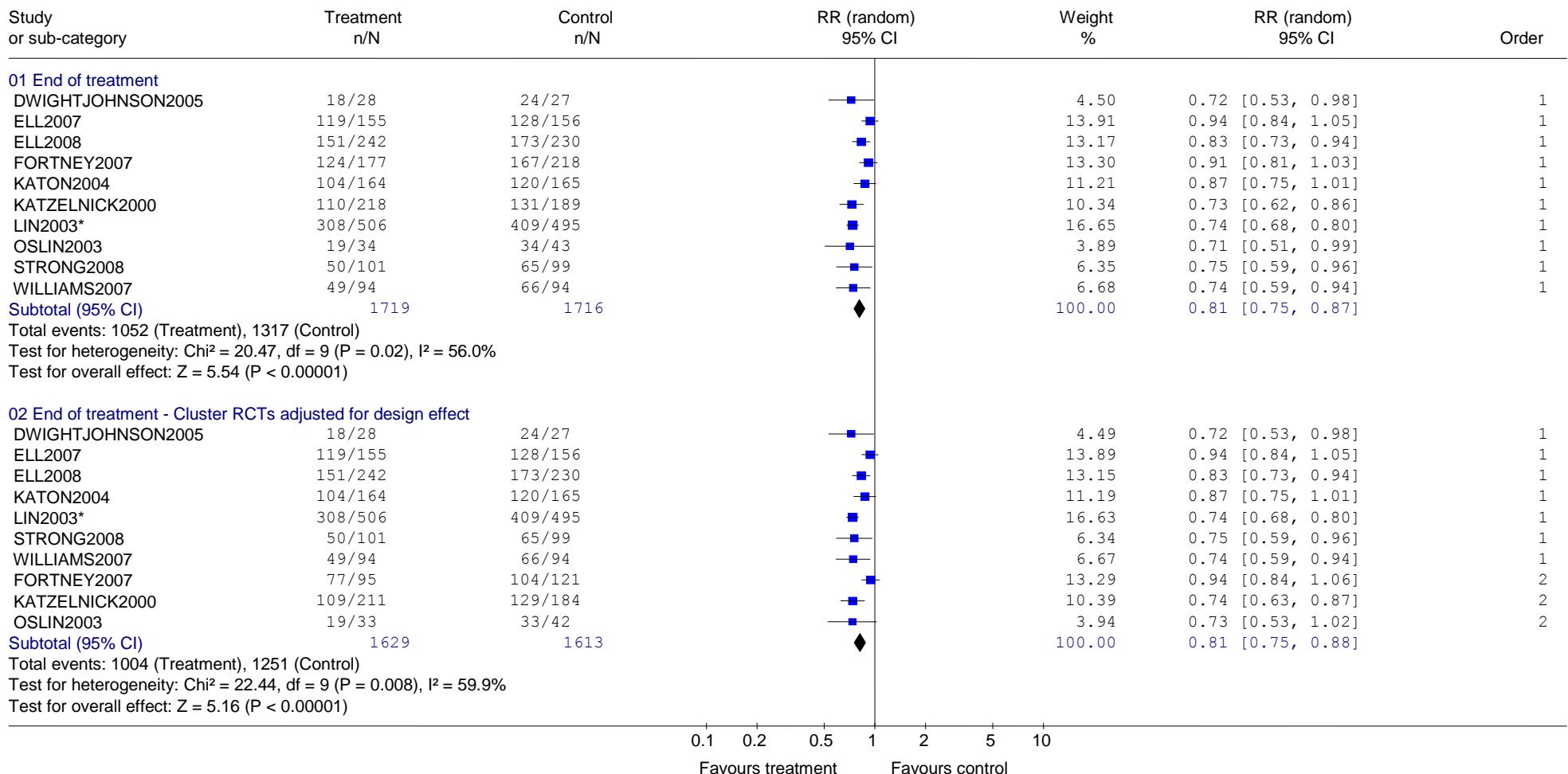


## Collaborative care – sensitivity analysis: ICC calculations

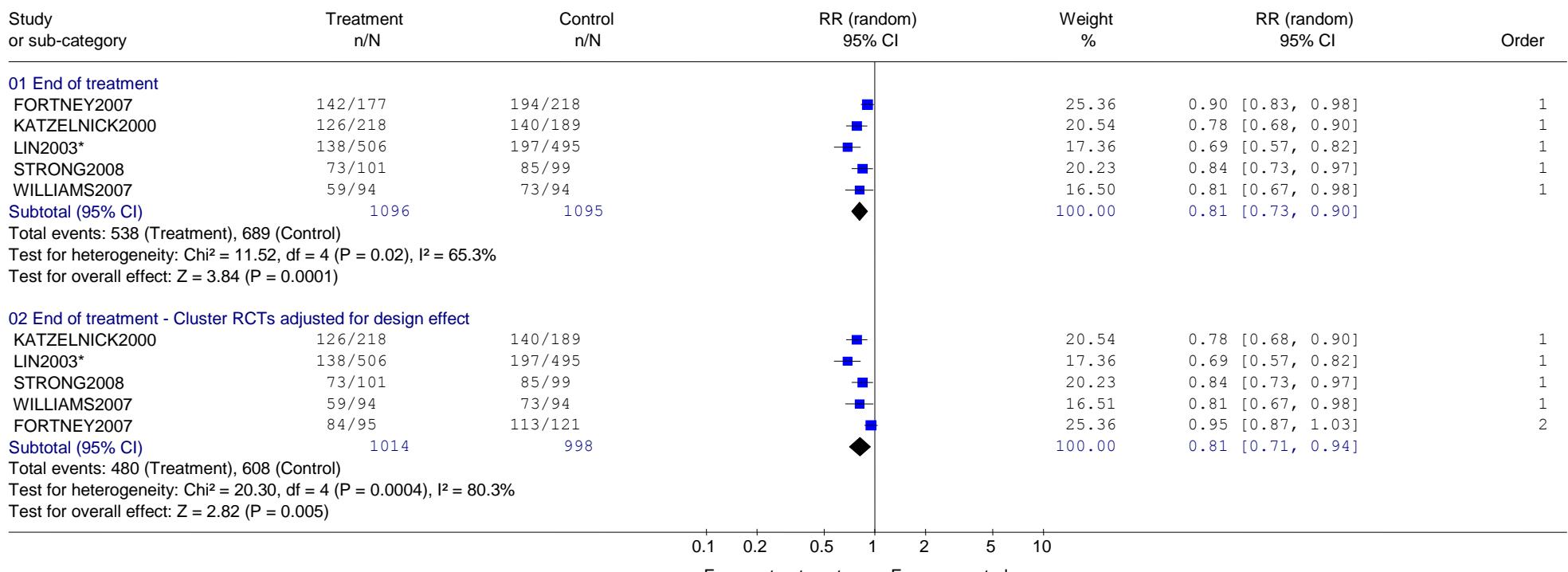
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 20 Mortality



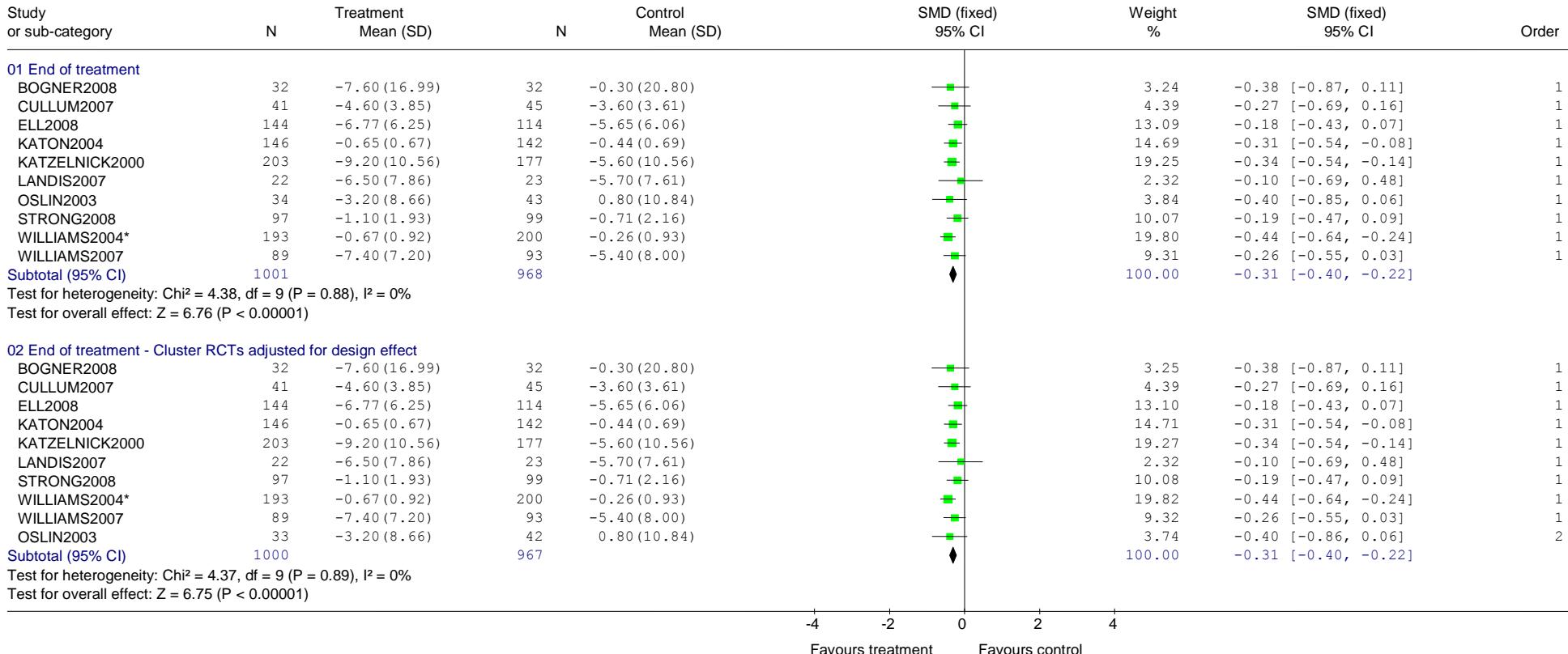
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 30 Depression outcome: 1. Non-response (<50% improvement) - outlier removed



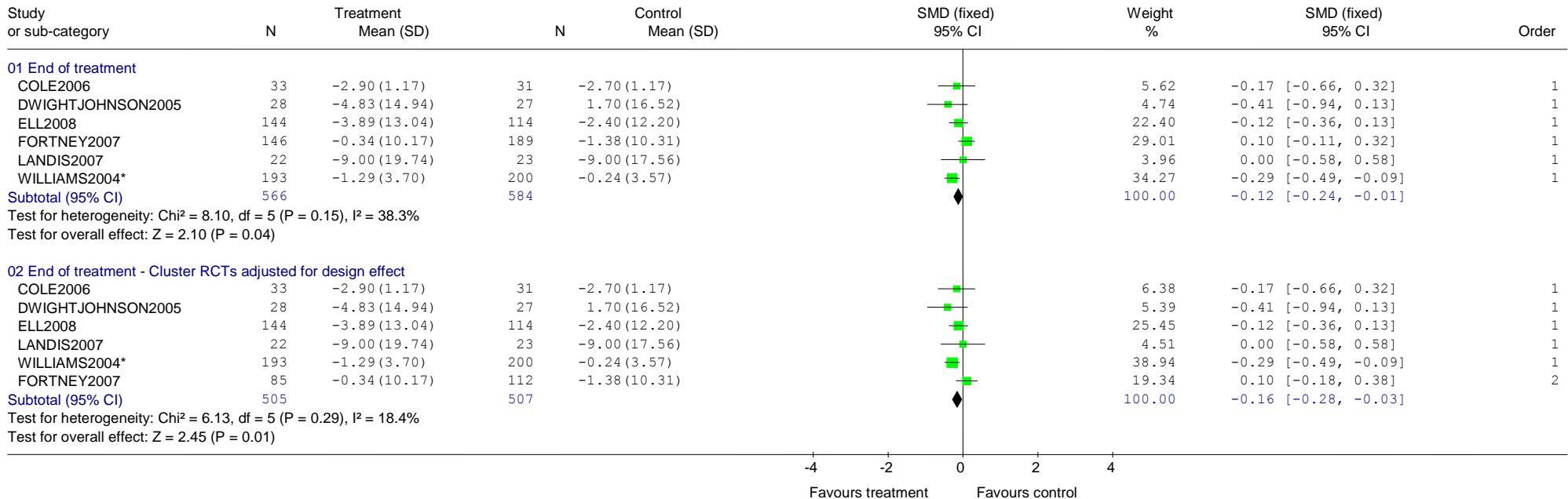
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 32 Depression outcome 2. Non-remission (scoring above cut-off for example, >7 on HAM-D) - outlier removed



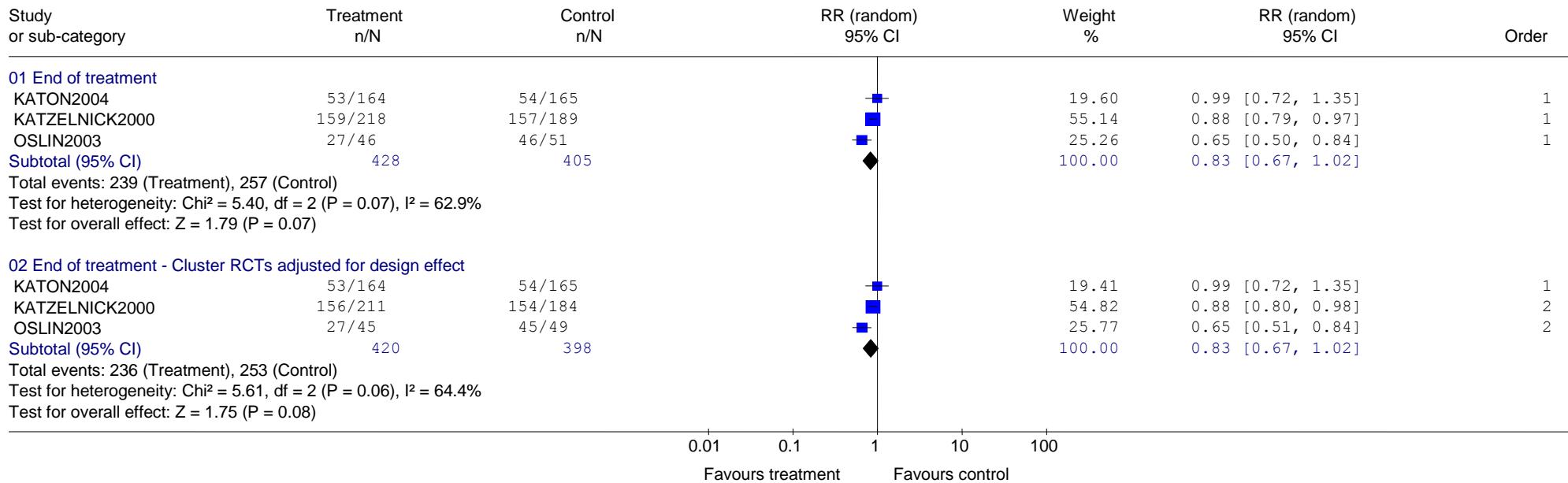
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 34 Depression outcome: 4.Continuous measures depression rating scale (Change score)



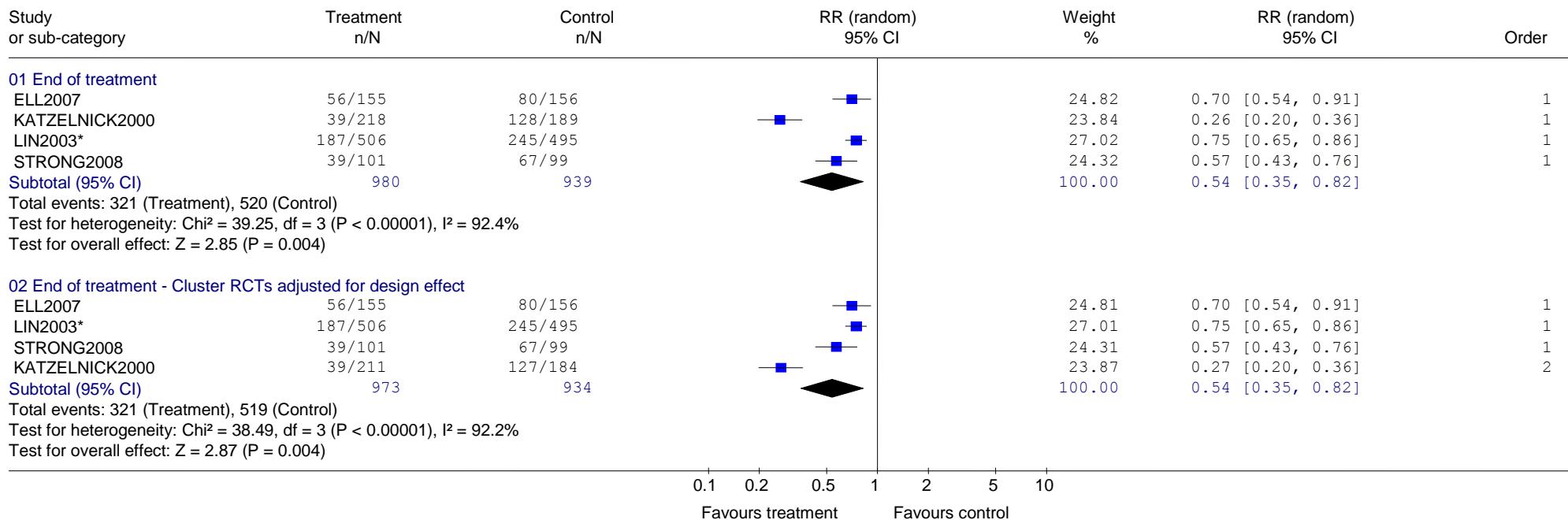
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 49 Physical health outcome/ QoL: 2. General physical well-being/ functioning (Change scores)



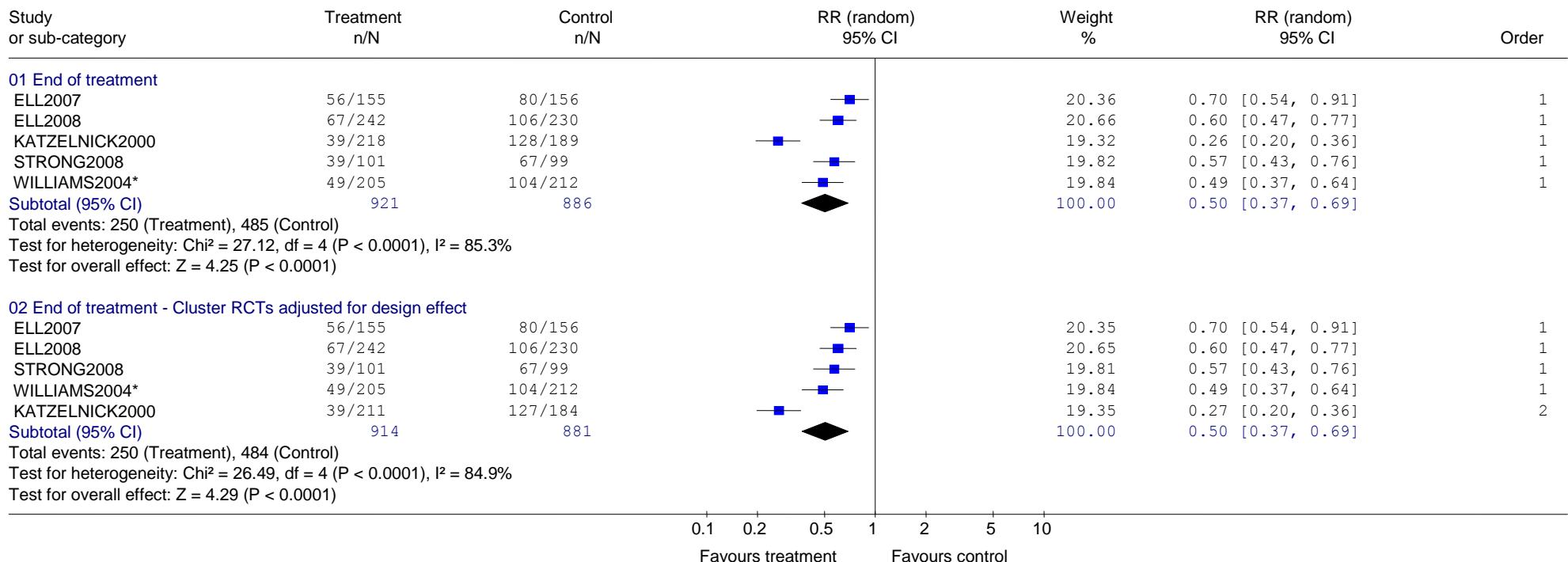
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything ( WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 60 Service use/ Process of care: 1. Did not receive a consultation / specified number of mental health visits



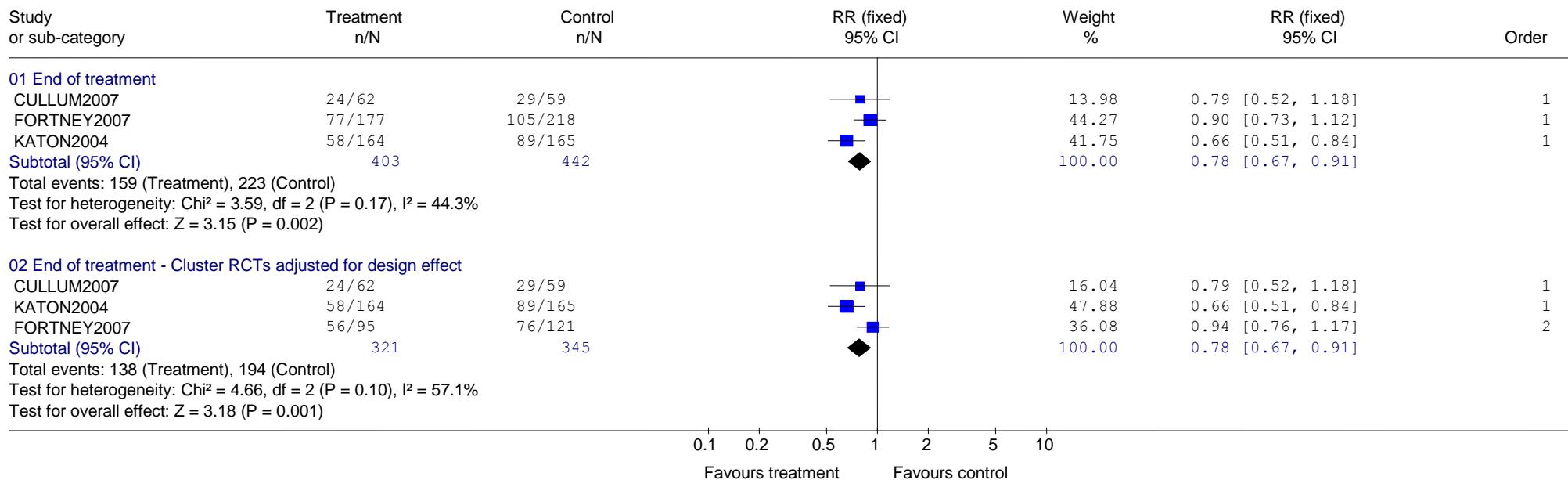
Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 65 Service use/ process of care : 6. Not receiving antidepressant medication



Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 67 Service use / Process of care (64-66 combined)



Review: DCHP - Service  
 Comparison: 03 Collaborative care versus everything (WCS analysis) - sensitivity analysis based on ICC  
 Outcome: 80 Satisfaction with service - not satisfied with treatment/care

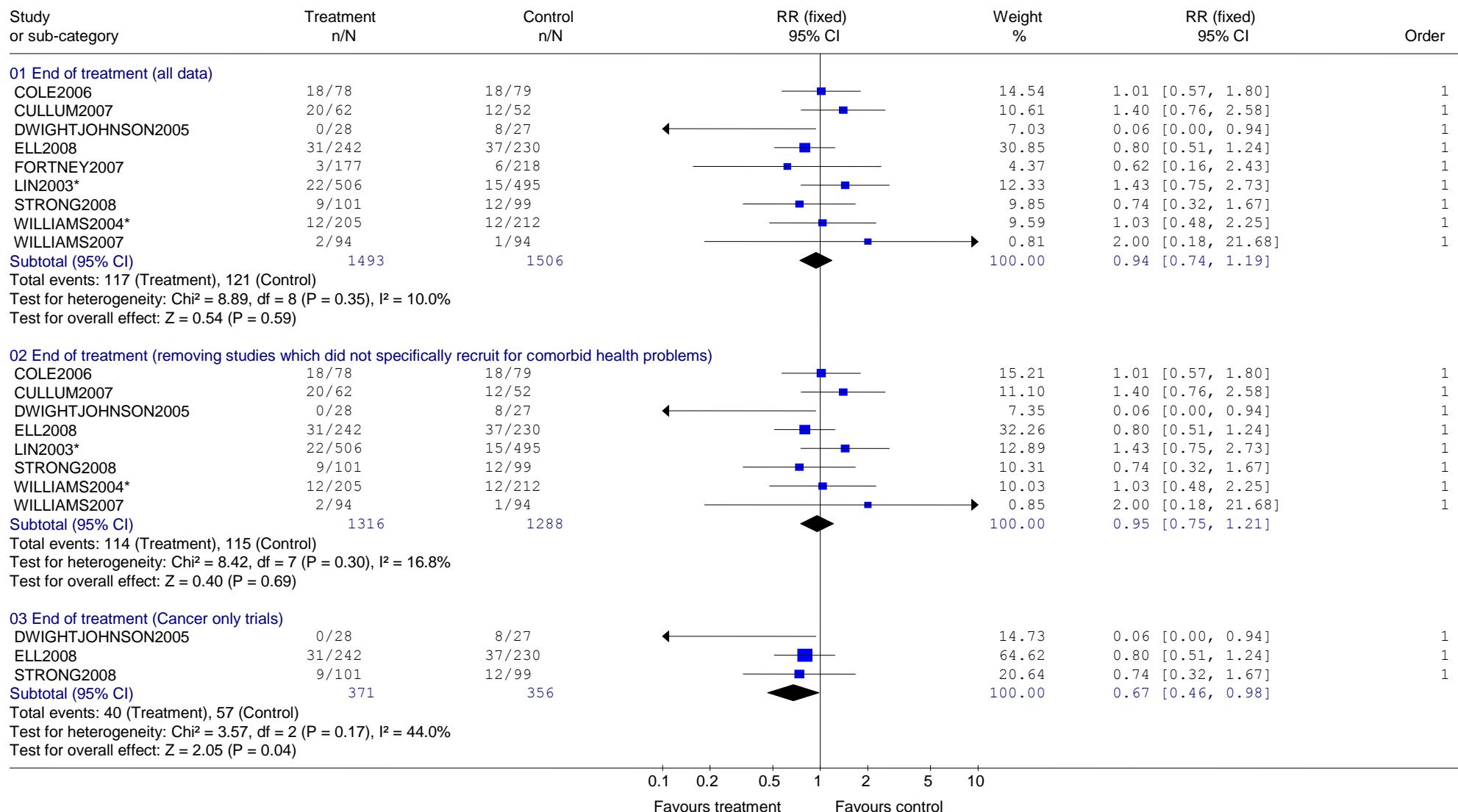


## Collaborative care – sensitivity analysis: recruited for comorbid physical health condition

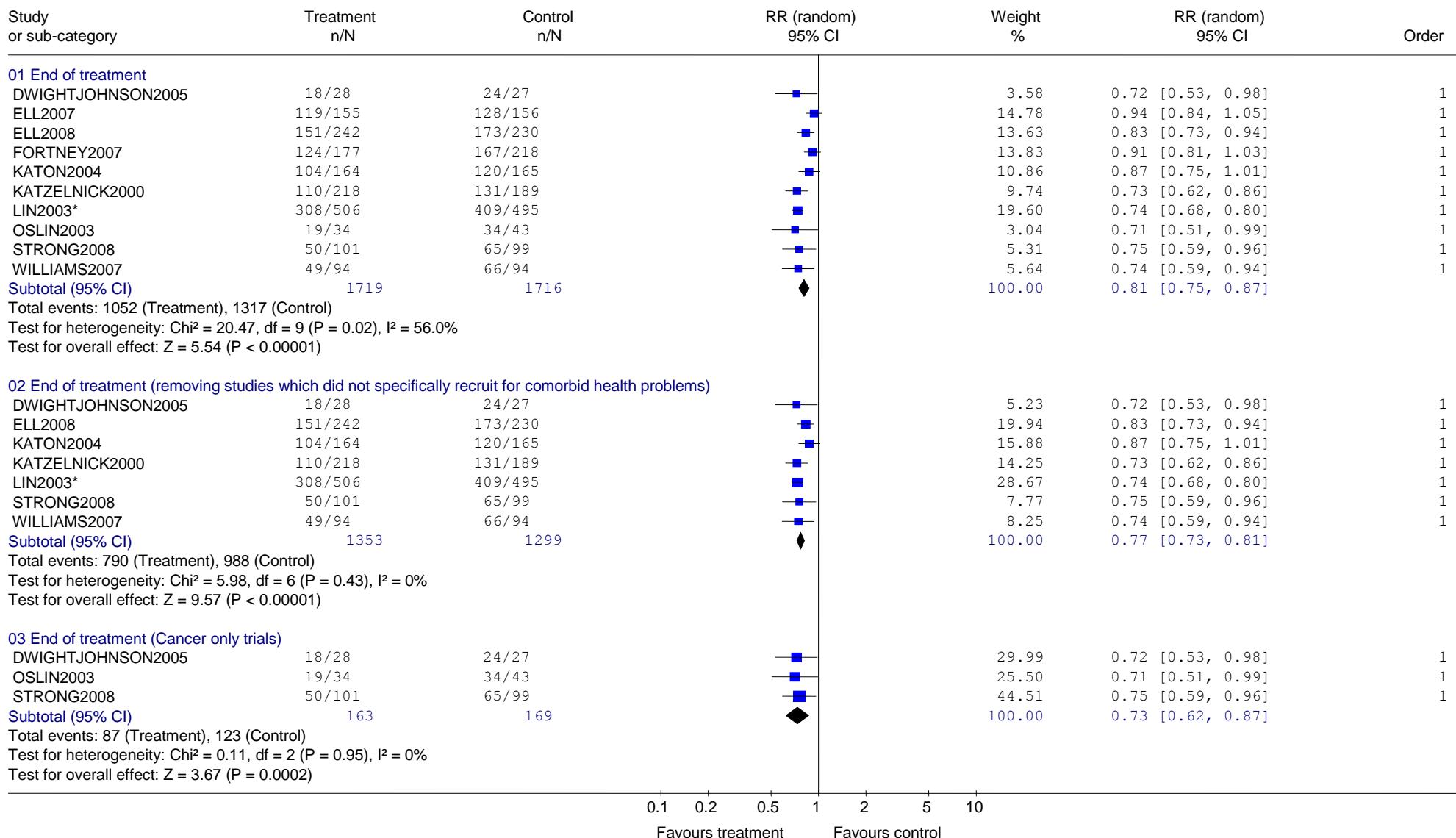
Review: DCHP - Service

Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)

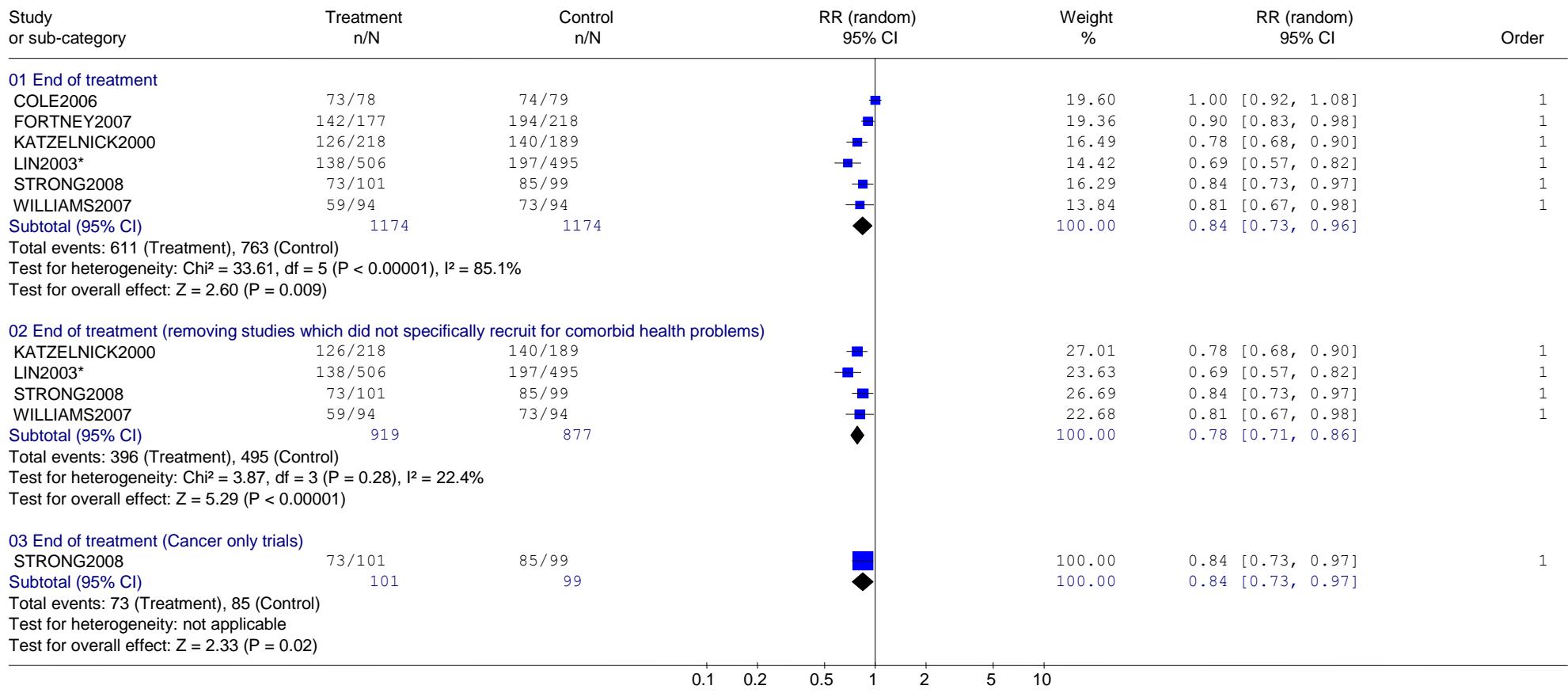
Outcome: 20 Mortality



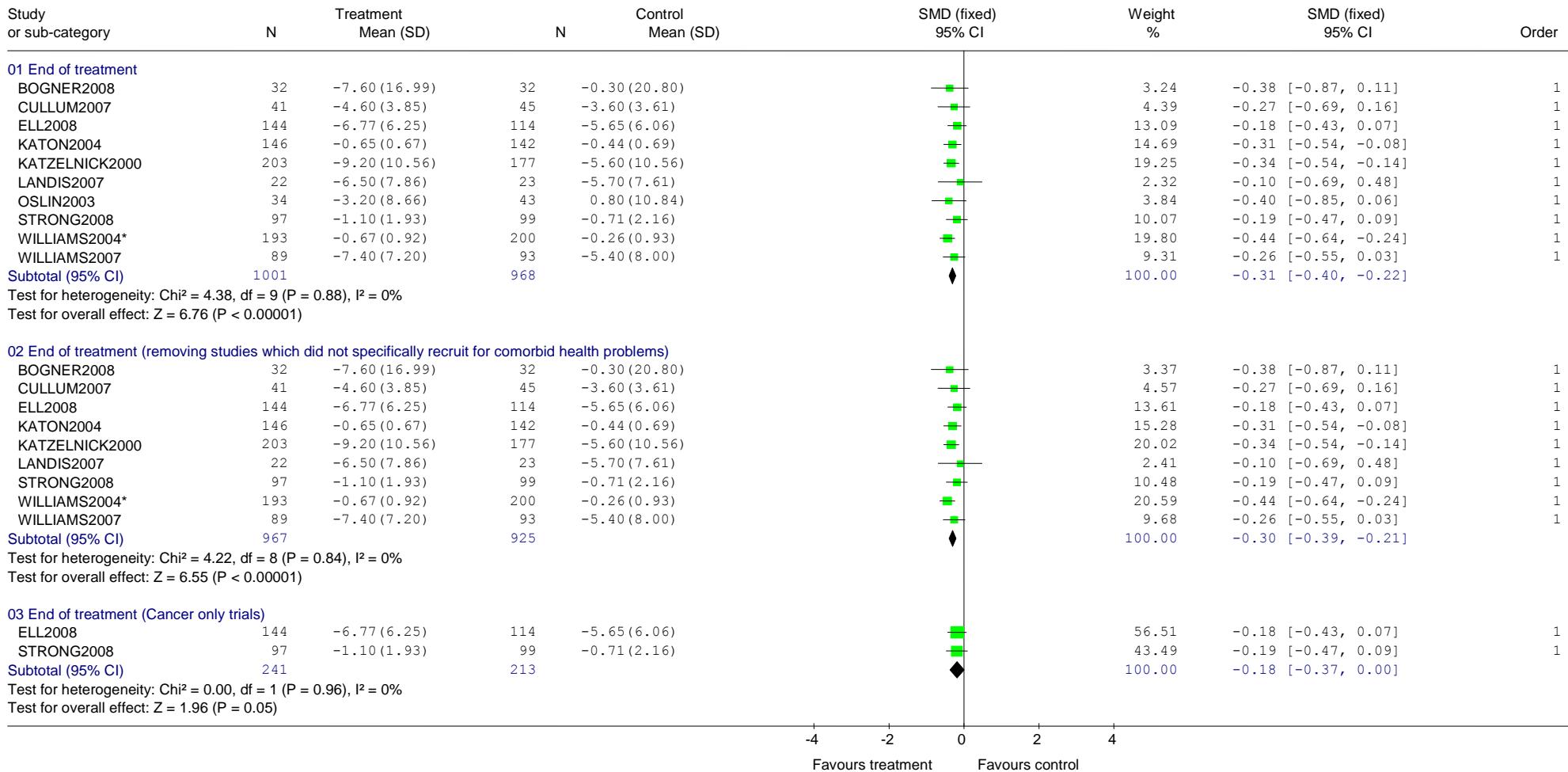
Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 30 Depression outcome: 1. Non-response (<50% improvement) - sensitivity analysis



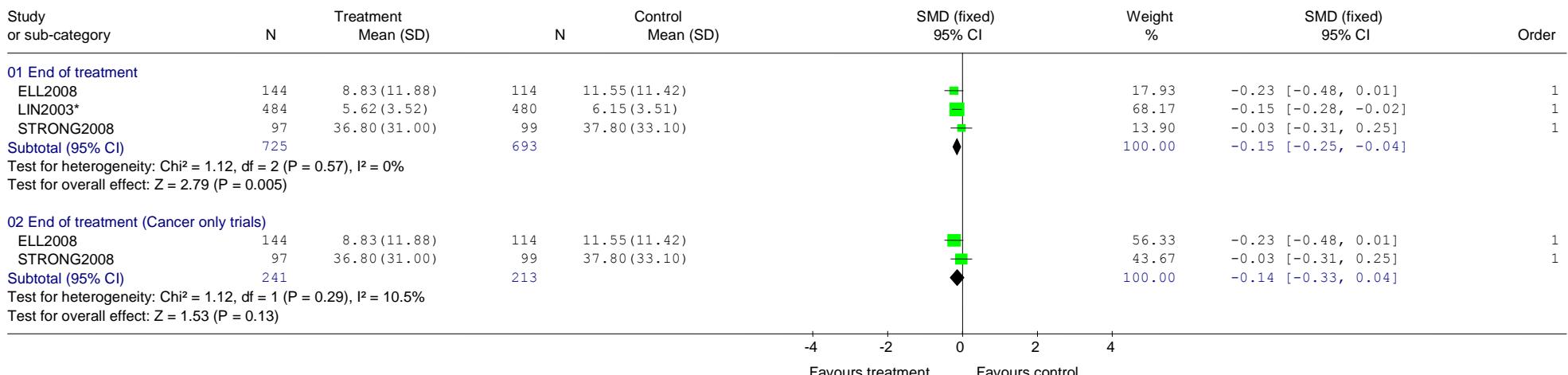
Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 32 Depression outcome: 2. Non-remission (scoring above cut off e.g. >7 on HAM-D)



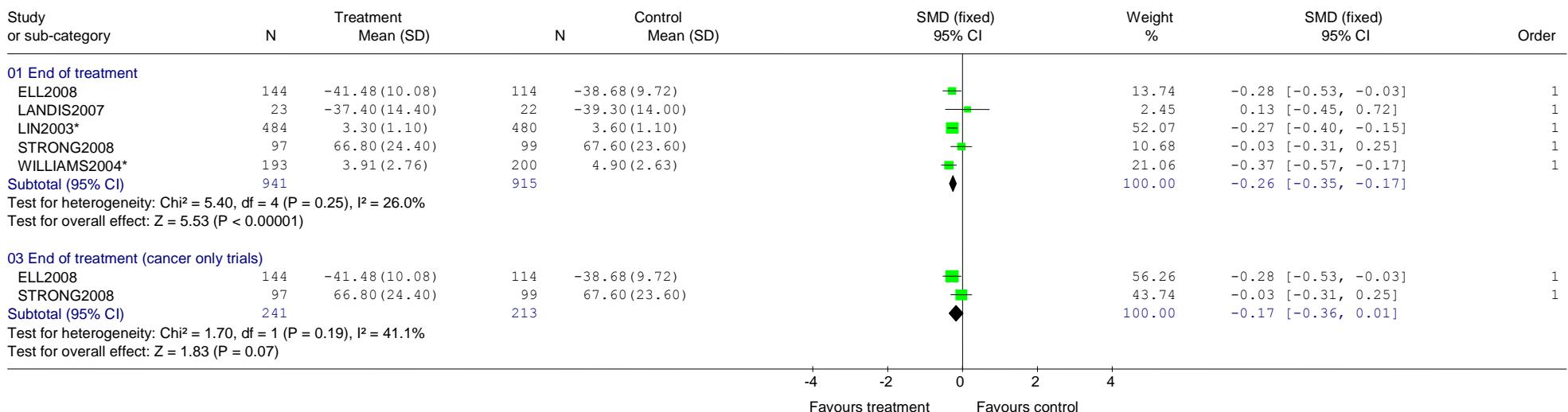
Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 34 Depression outcome: 4.Continuous measures depression rating scale (Change score)



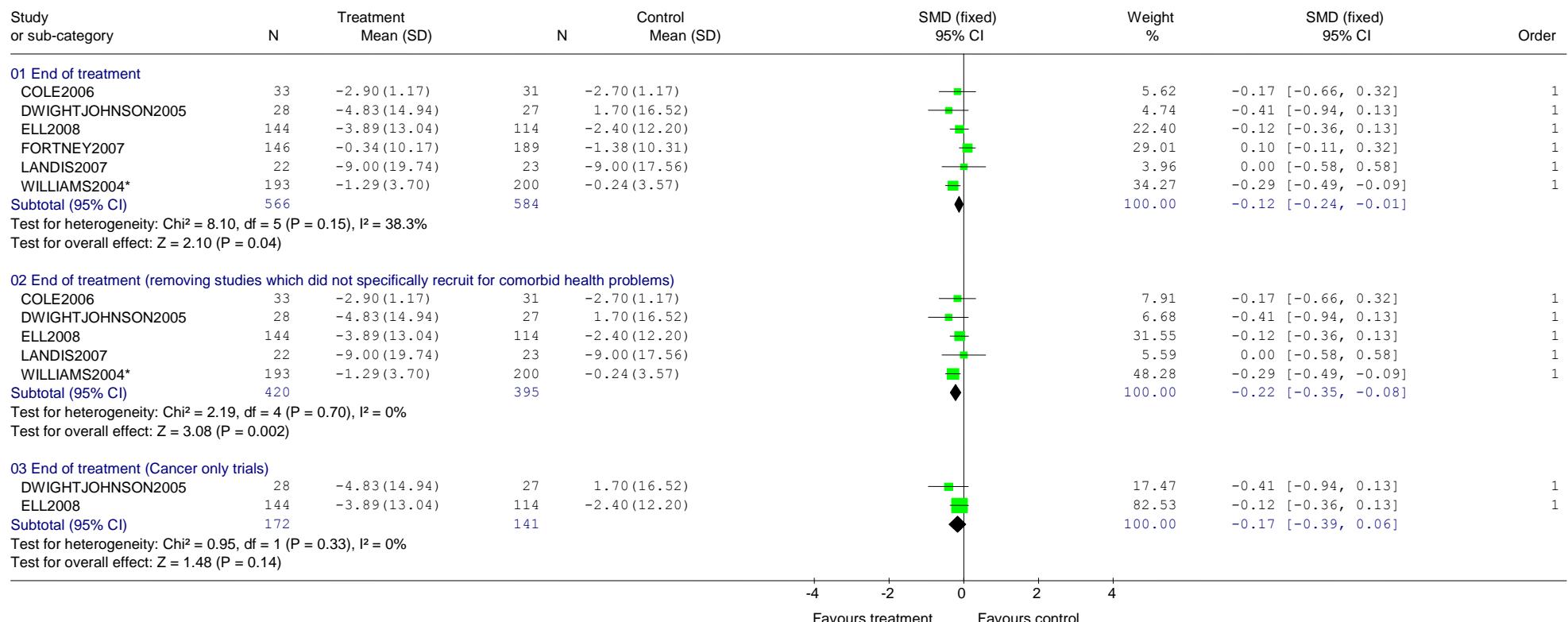
Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 41 Physical health outcome: 2. Pain intensity (Brief Pain Inventory, author defined scale)



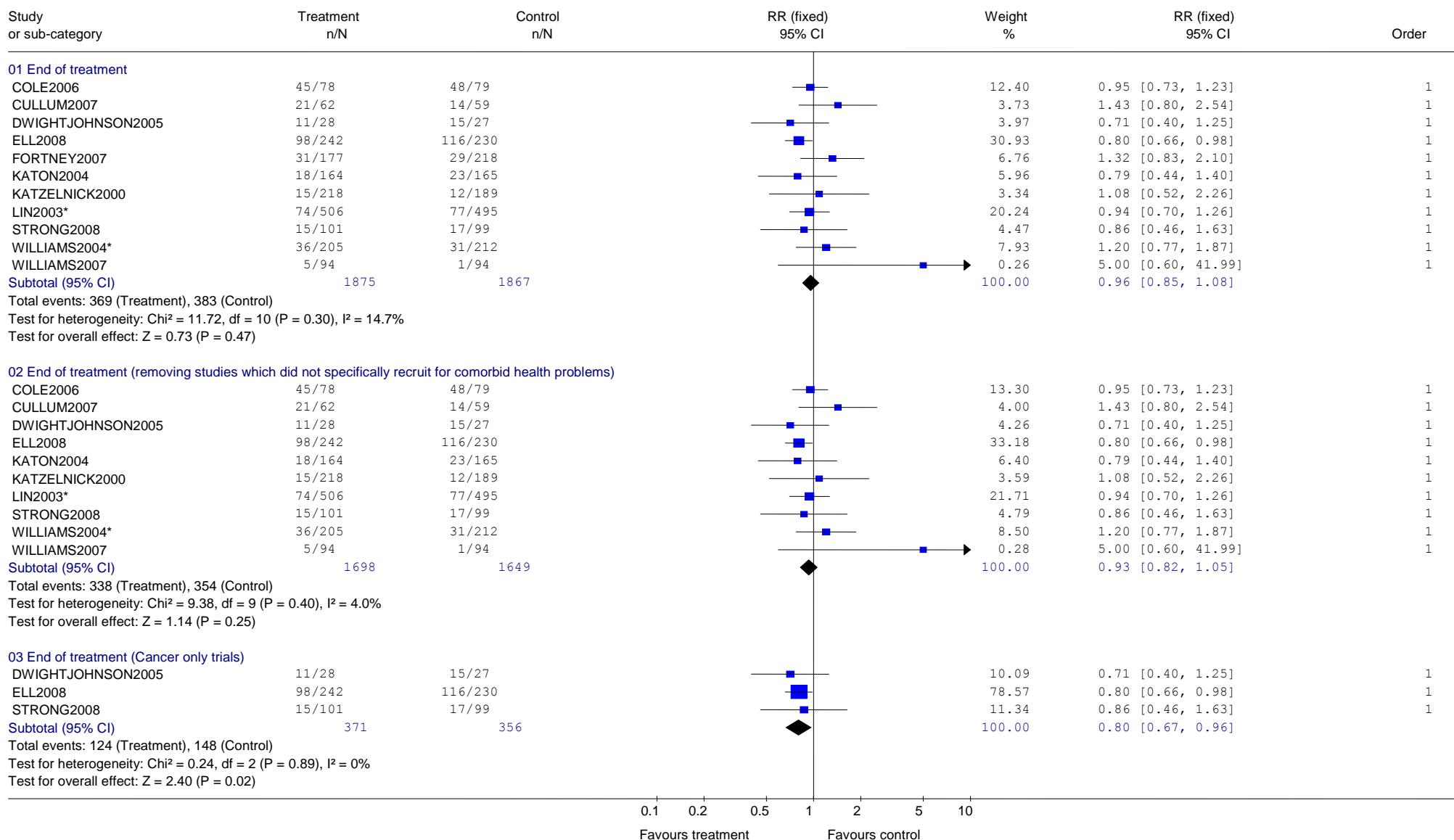
Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 49 Physical health outcome/ QoL: 1. General physical well-being/ functioning (SF-12 physical etc)



Review: DCHP - Service  
 Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
 Outcome: 50 Physical health outcome/ QoL: 2. General physical well-being/ functioning (Change scores)

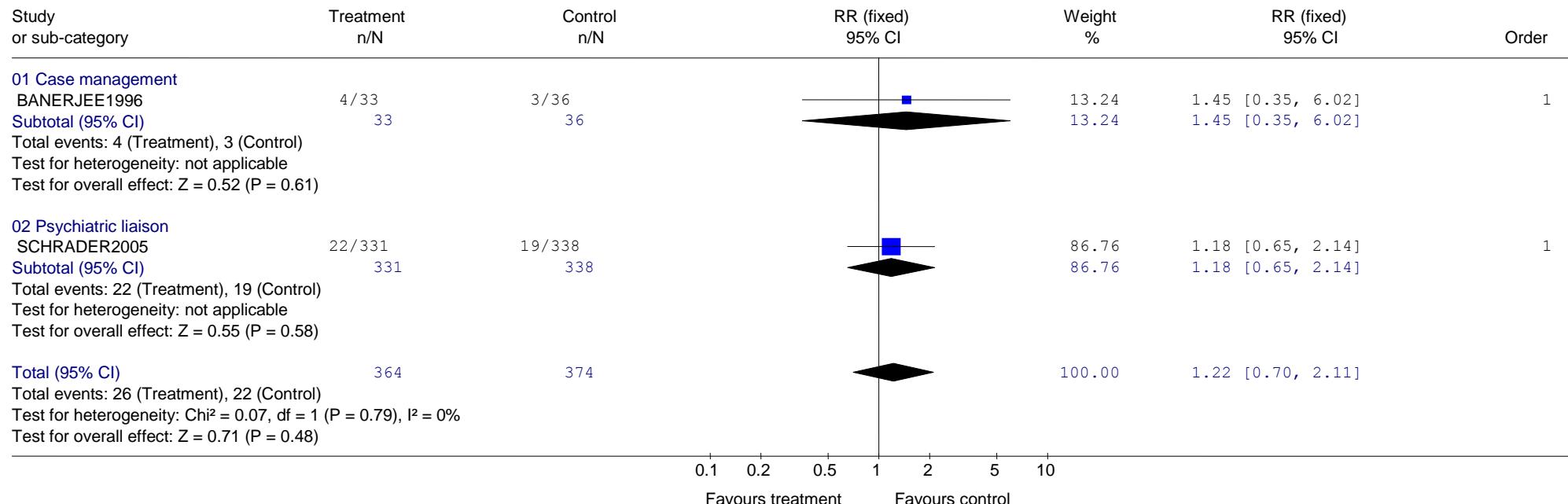


Review: DCHP - Service  
Comparison: 05 Collaborative care versus everything (WCS analysis) - sensitivity analysis (recruited for chronic condition)  
Outcome: 70 Treatment acceptability - leaving the study early for any reason

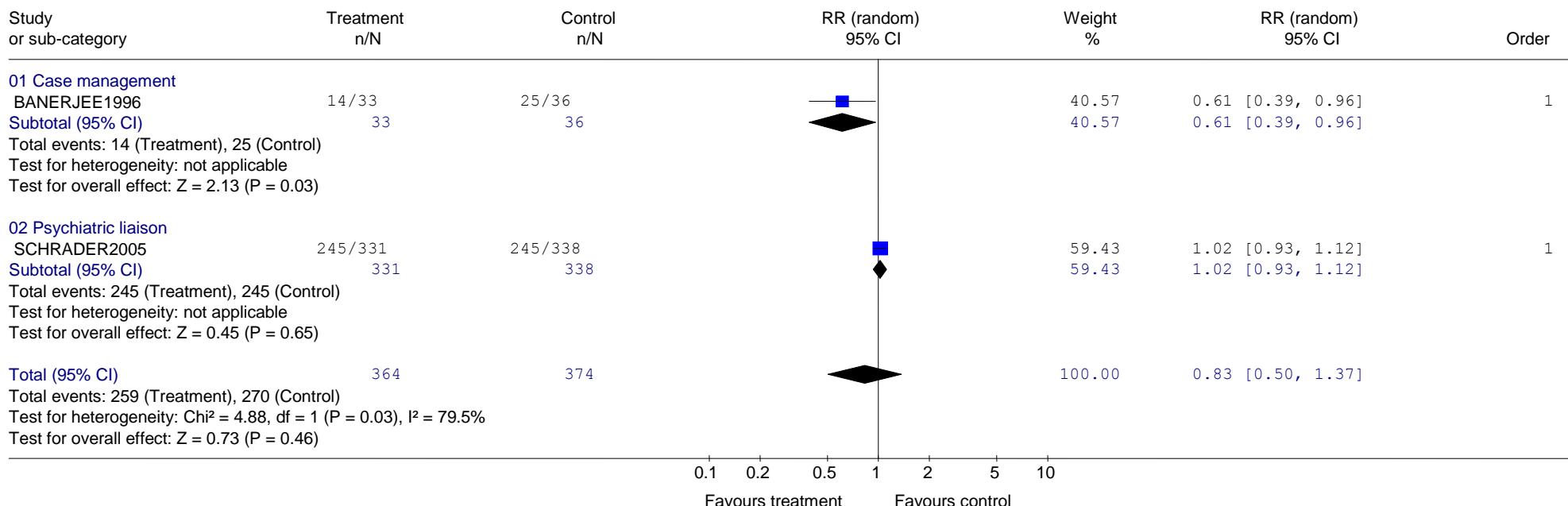


## Other service-level interventions versus standard care

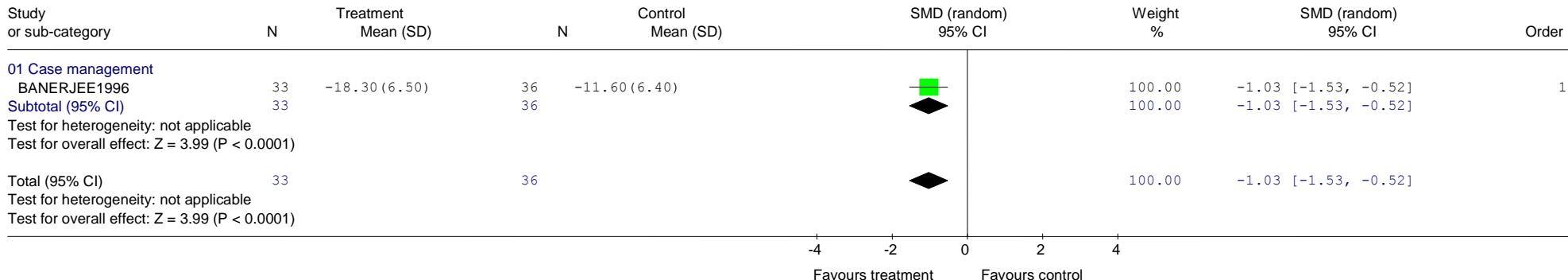
Review: DCHP - Service  
 Comparison: 20 Other service-level interventions (multidisciplinary teams, consultation-liaison) versus standard care (WCS)  
 Outcome: 20 Mortality



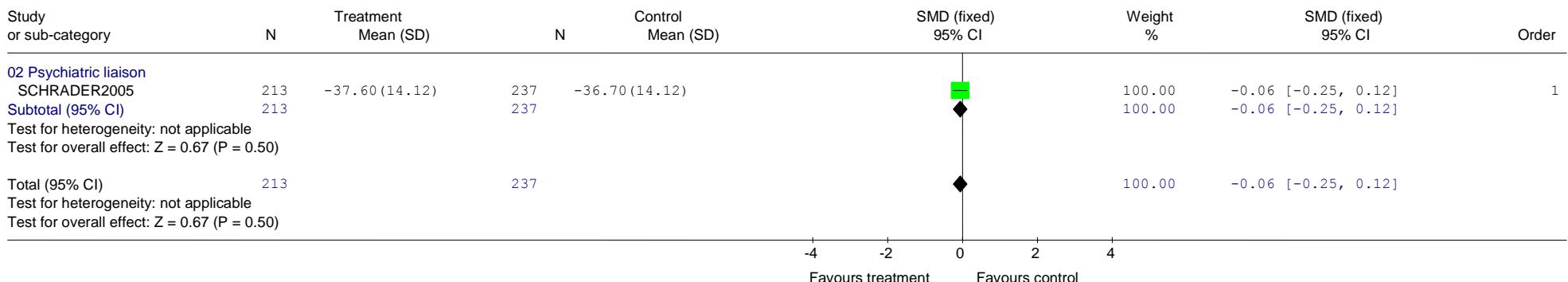
Review: DCHP - Service  
 Comparison: 20 Other service level interventions (multidisciplinary teams, consultation-liaison) versus standard care (WCS)  
 Outcome: 33 Depression outcome: 3. Diagnosis (at follow-up)



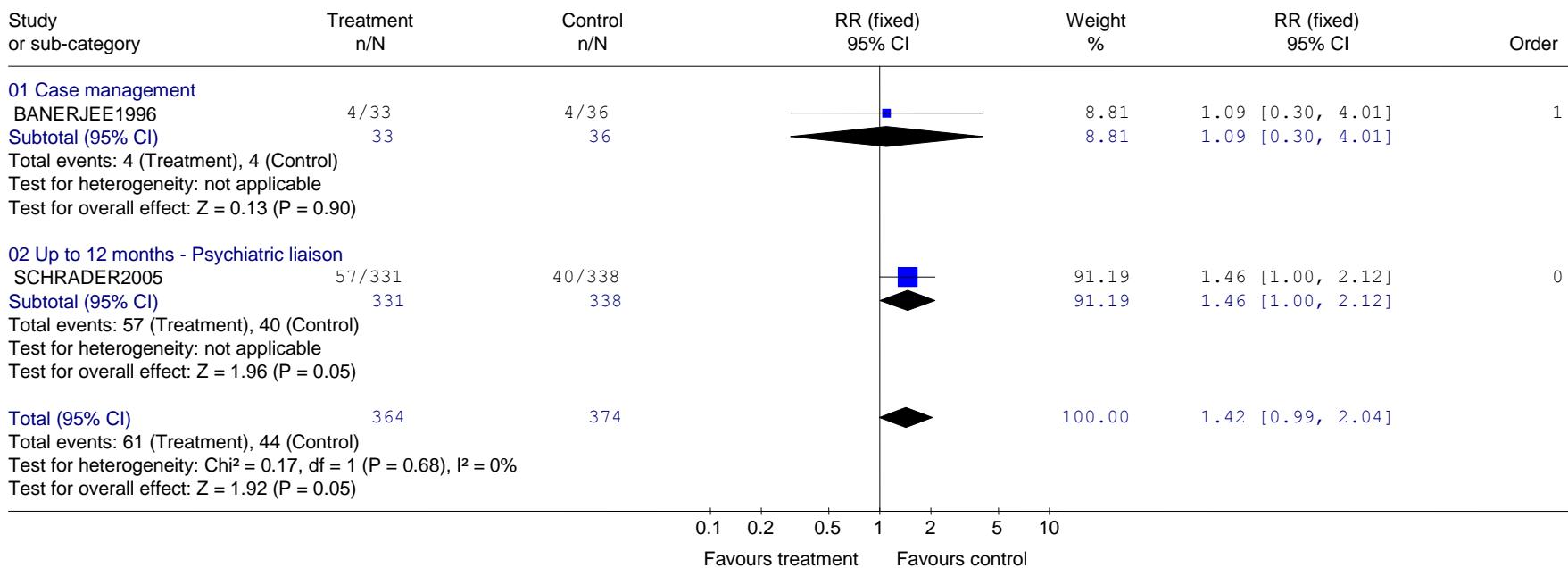
Review: DCHP - Service  
 Comparison: 20 Other service level interventions (multidisciplinary teams, consultation-liaison) versus standard care (WCS)  
 Outcome: 34 Depression outcome: 4.Continuous measures depression rating scale (Change score)



Review: DCHP - Service  
 Comparison: 20 Other service level interventions (multidisciplinary teams, consultation-liaison) versus standard care (WCS)  
 Outcome: 48 Physical health outcome/ QoL: 1. General physical well-being/ functioning (SF-12 physical etc)



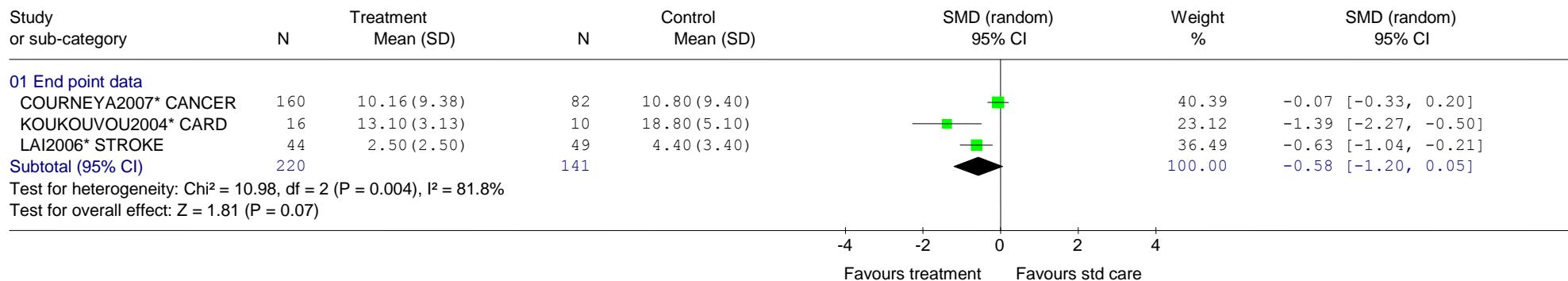
Review: DCHP - Service  
 Comparison: 20 Other service level interventions (multidisciplinary teams, consultation-liaison) versus standard care (WCS)  
 Outcome: 70 Treatment acceptability - leaving the study early for any reason



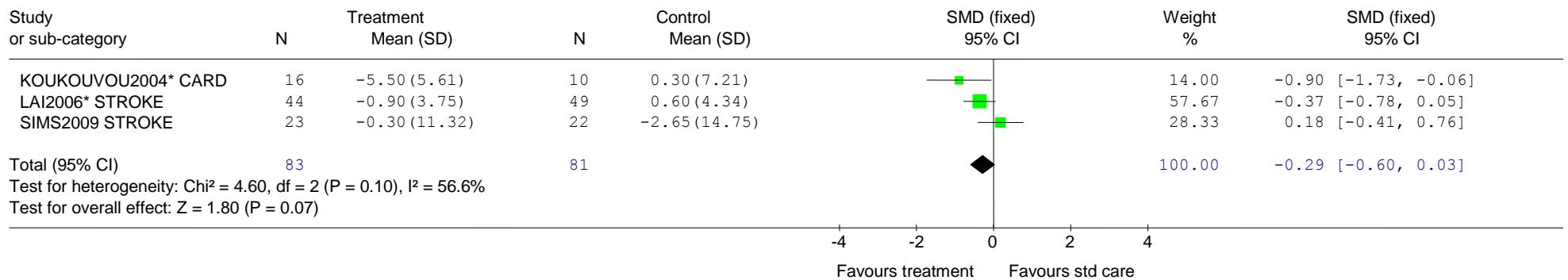
## *Psychological and psychosocial interventions*

### **Physical activity versus standard care**

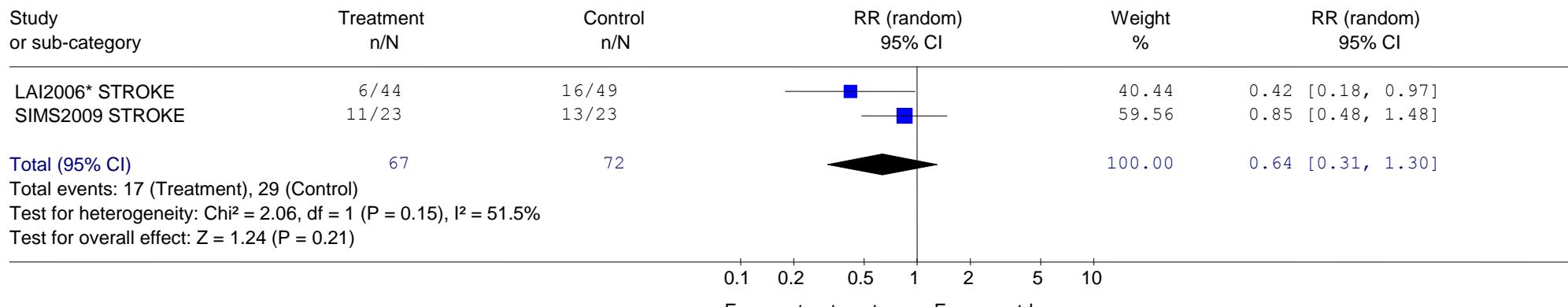
Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard (std) care  
 Outcome: 01 Depression (end of treatment)



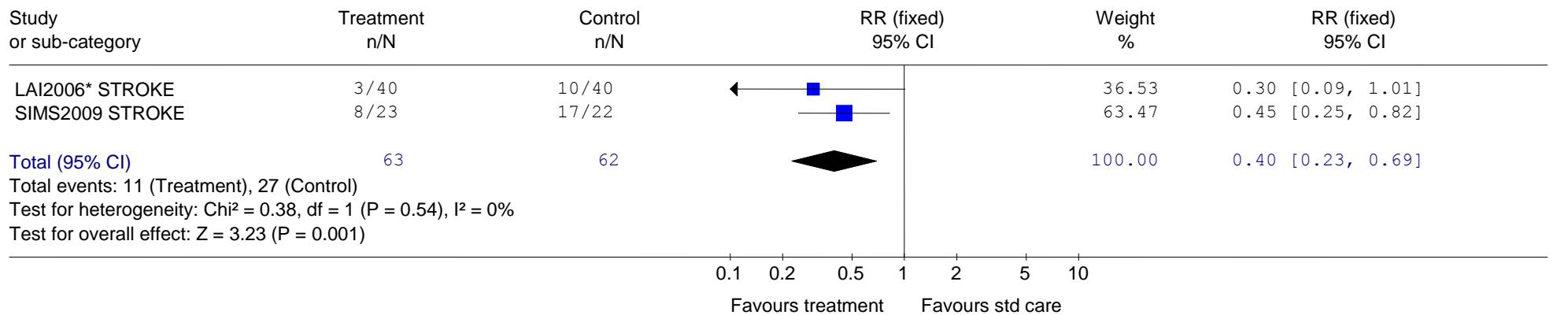
Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard care  
 Outcome: 02 Depression (change score: end of treatment)



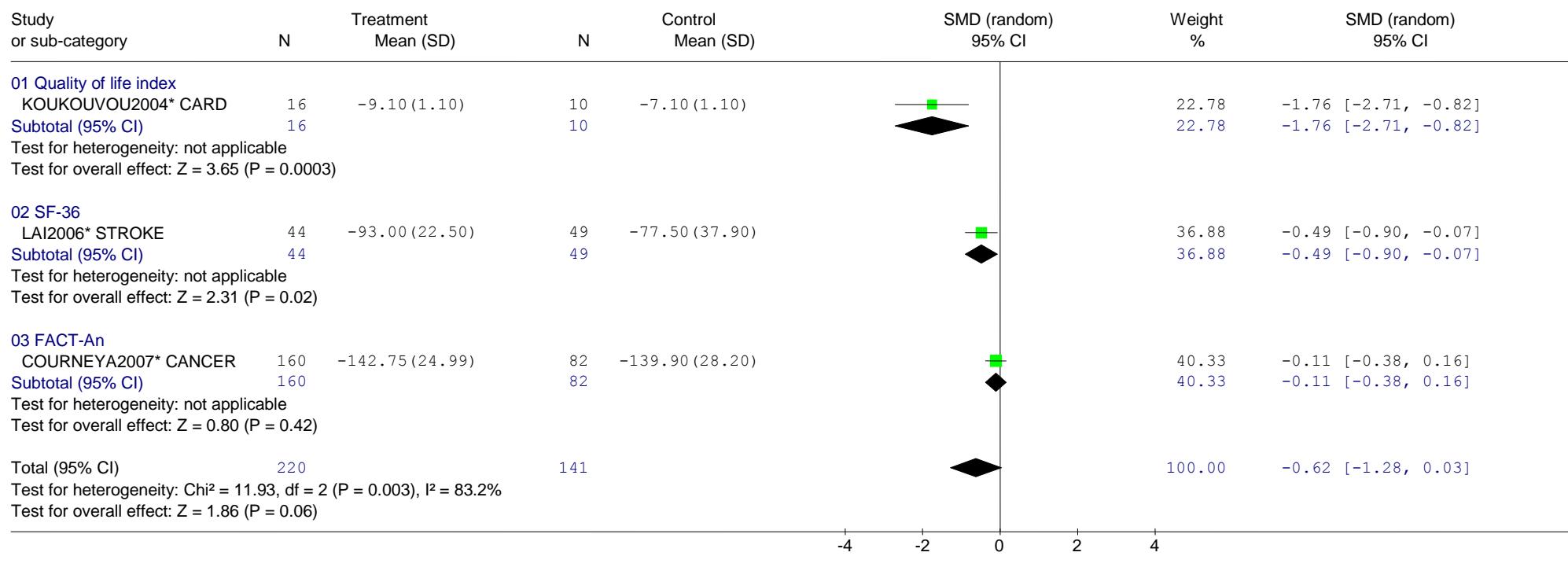
Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard care  
 Outcome: 03 Non-remission (below cut-off)



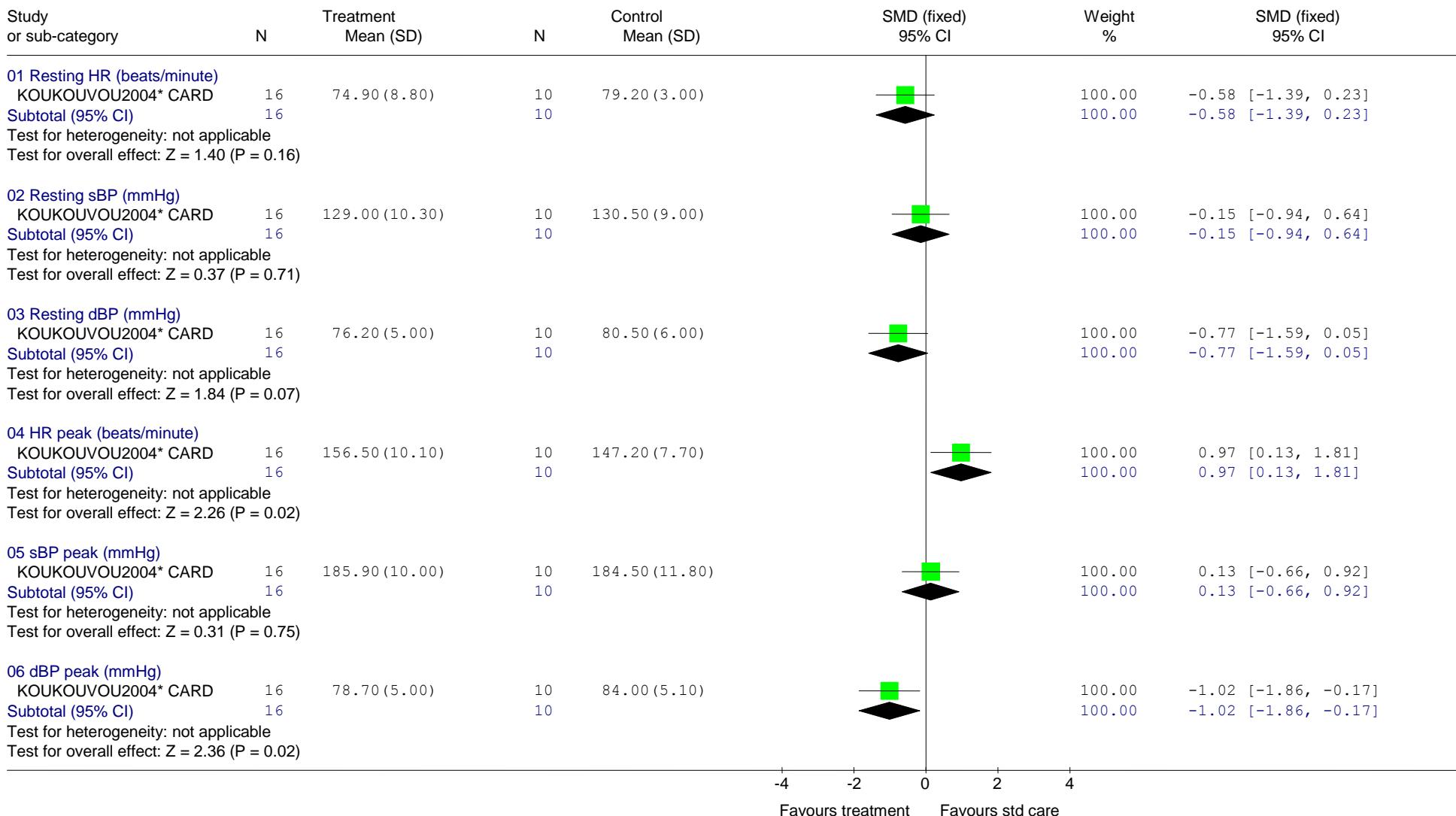
Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard care  
 Outcome: 04 Non-remission (6-month follow-up)



Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard care  
 Outcome: 07 Quality of life (end of treatment)



Review: DCHP: Physical activity  
 Comparison: 01 Physical activity versus standard care  
 Outcome: 08 Physical health outcomes (end of treatment)

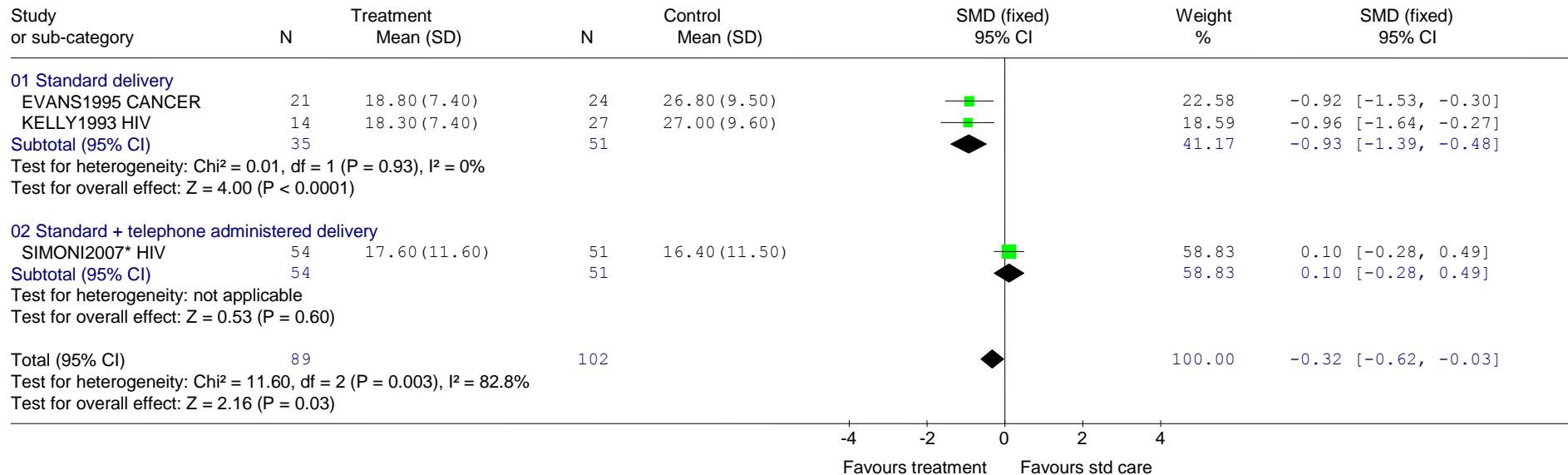


## Peer (self-help) support versus standard care

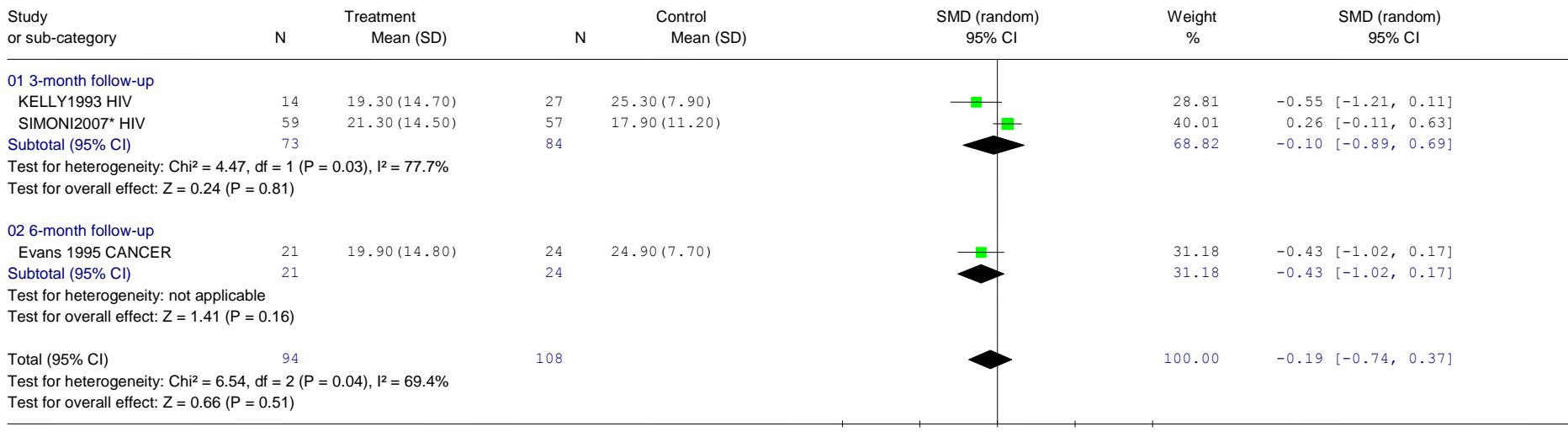
Review: DCHP: Peer (self-help) support

Comparison: 01 Peer (self-help) support versus standard care

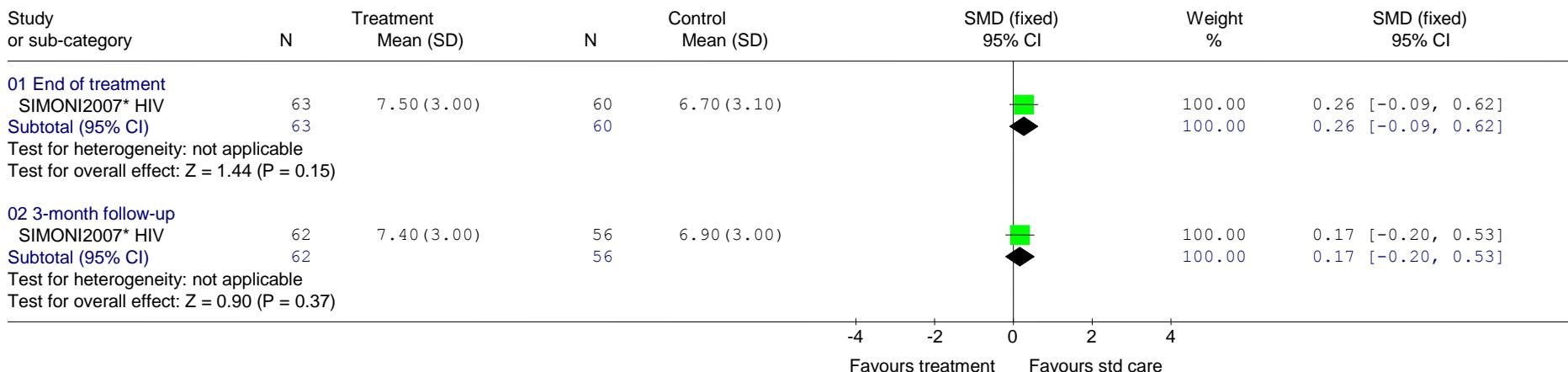
Outcome: 01 Depression (end of treatment)



Review: DCHP: Peer (self-help) support  
 Comparison: 01 Peer (self-help) support versus standard care  
 Outcome: 03 CES-D (6-month follow-up)



Review: DCHP: Peer (self-help) support  
 Comparison: 01 Peer (self-help) support versus standard care  
 Outcome: 04 Physical Health Outcomes: HIV-1 RNA viral load

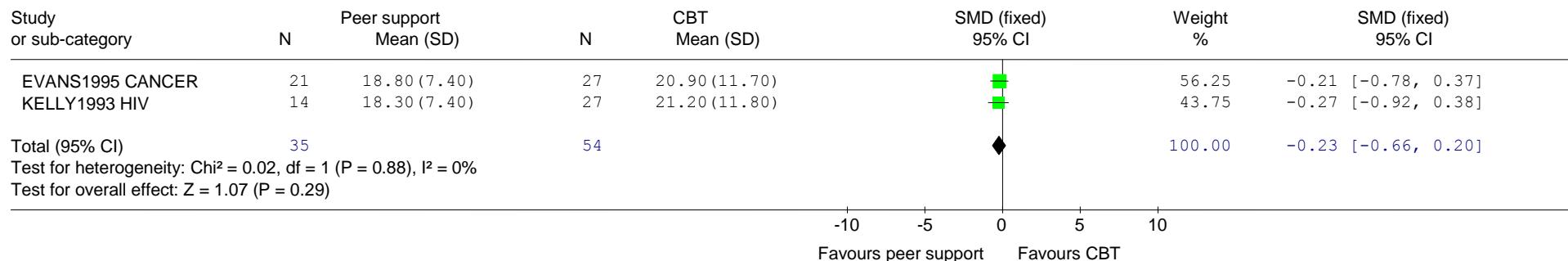


## Peer (self-help) support versus group based cognitive and behavioural interventions

Review: DCHP: Peer (self-help) support

Comparison: 02 Peer (self-help) support versus group based cognitive and behavioural interventions

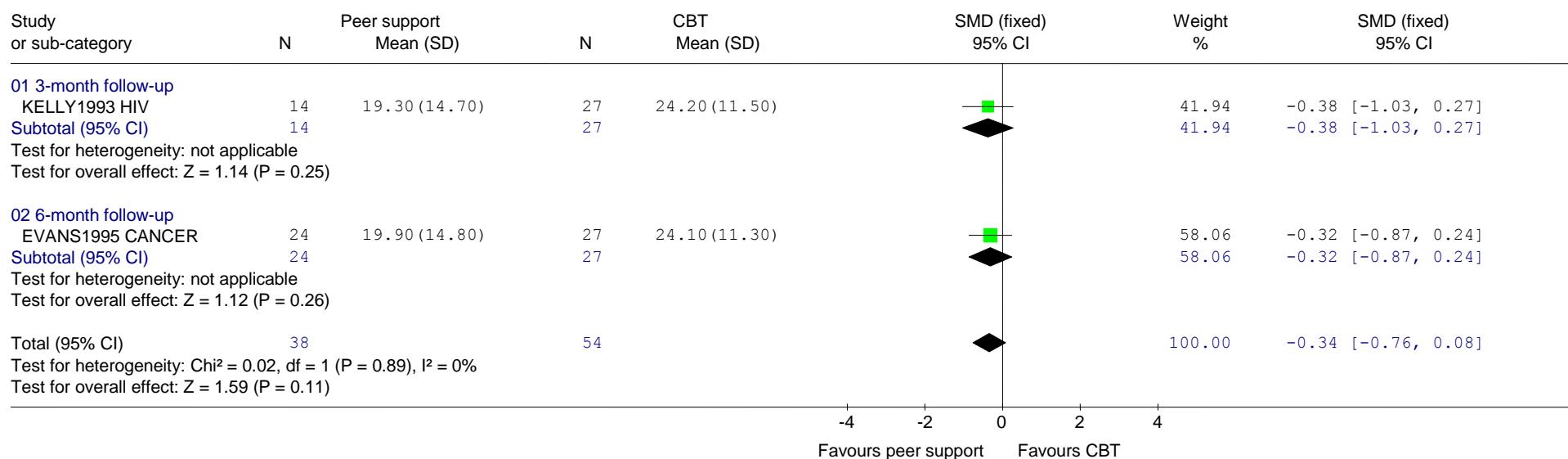
Outcome: 01 Depression (end of treatment)



Review: DCHP: Peer (self-help) support

Comparison: 02 Peer (self-help) support versus group based cognitive and behavioural interventions

Outcome: 02 Depression (follow-up)

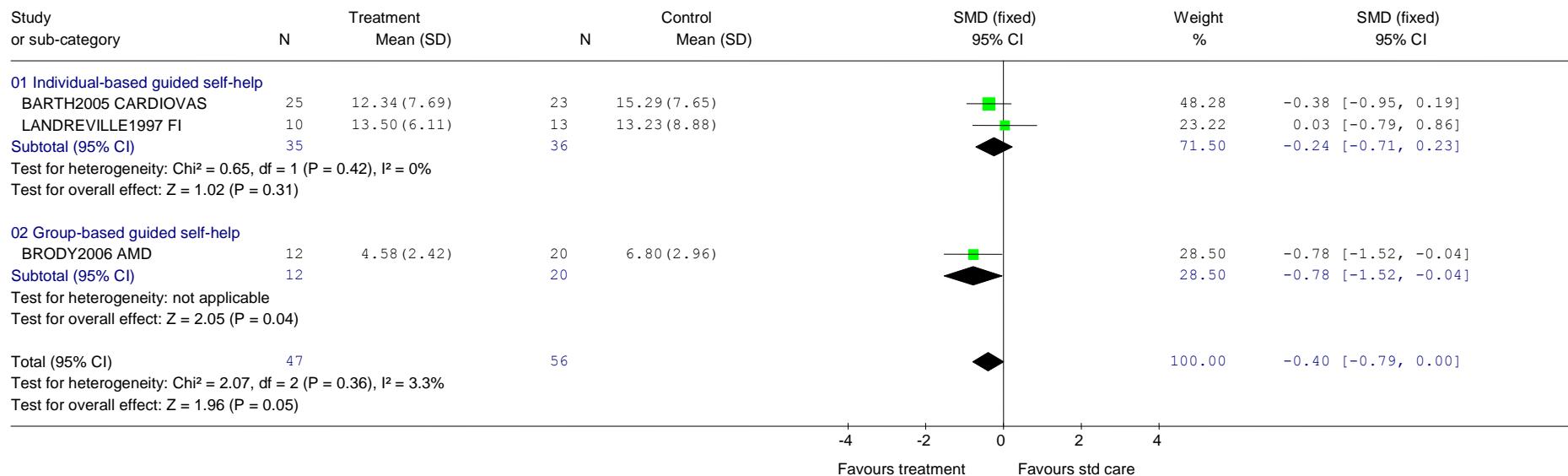


## Guided self-help based on cognitive and behavioural principles versus standard care

Review: DCHP: Guided self-help

Comparison: 01 Guided self-help based on cognitive behavioural principles versus standard care

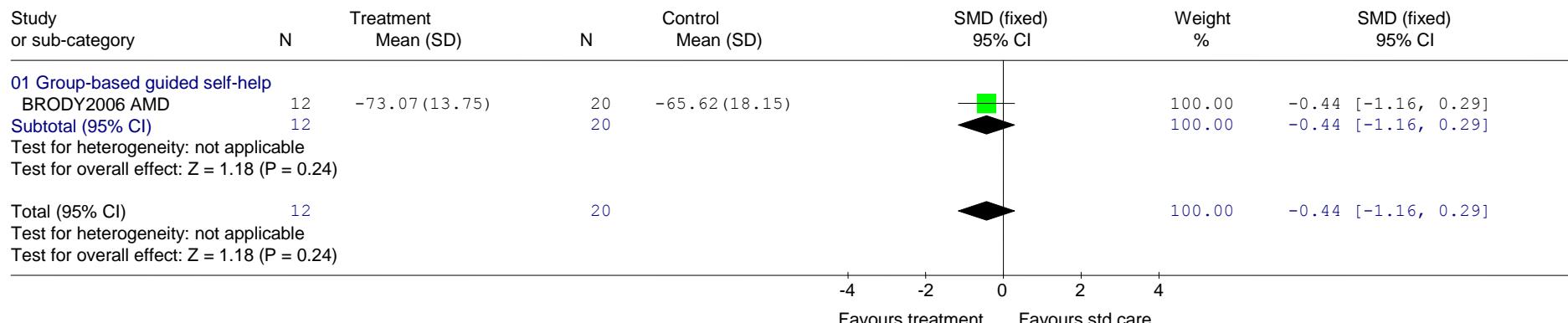
Outcome: 01 Depression (end of treatment)



Review: DCHP: Guided self-help (not separating comparison)

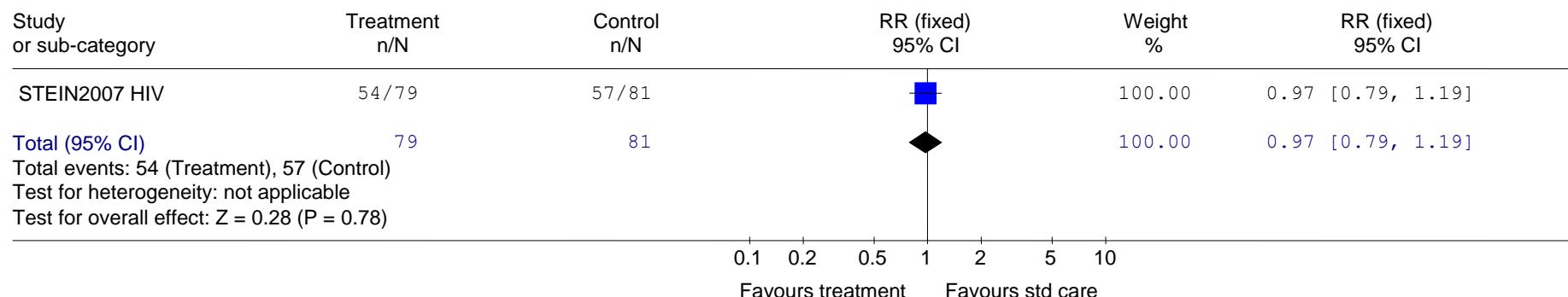
Comparison: 01 Guided self-help based on cognitive behavioural principles versus standard care

Outcome: 02 Physical health outcome: visual functioning questionnaire

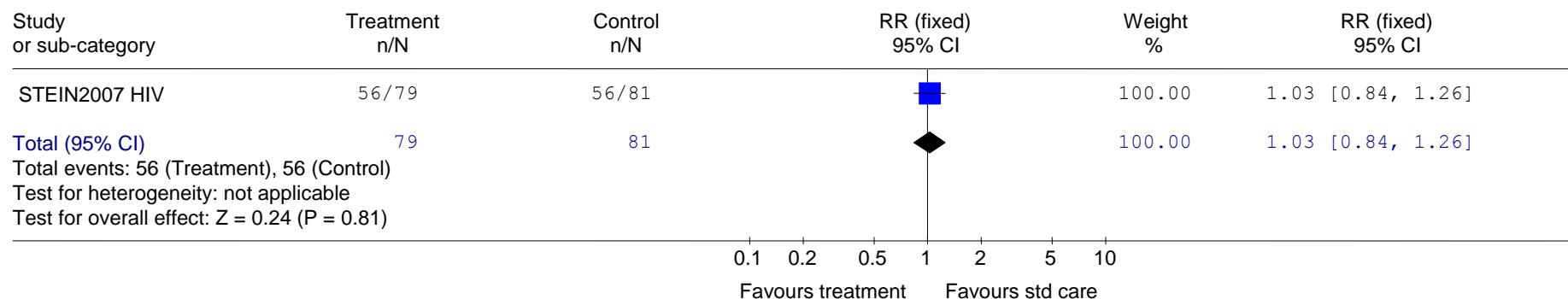


## Individual guided self-help based on McMaster model of family functioning versus standard care

Review: DCHP: Individual guided self-help  
 Comparison: 02 Individual guided self-help based on McMaster model of family functioning principles versus standard care  
 Outcome: 01 Non-remission (below cut-off)

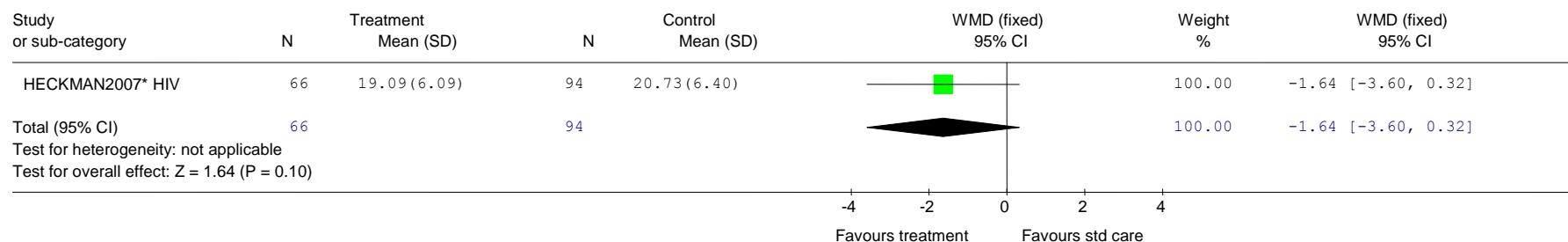


Review: DCHP: Individual guided self-help  
 Comparison: 02 Individual guided self-help based on McMaster model of family functioning principles versus standard care  
 Outcome: 02 Non-response (<50% reduction from baseline)

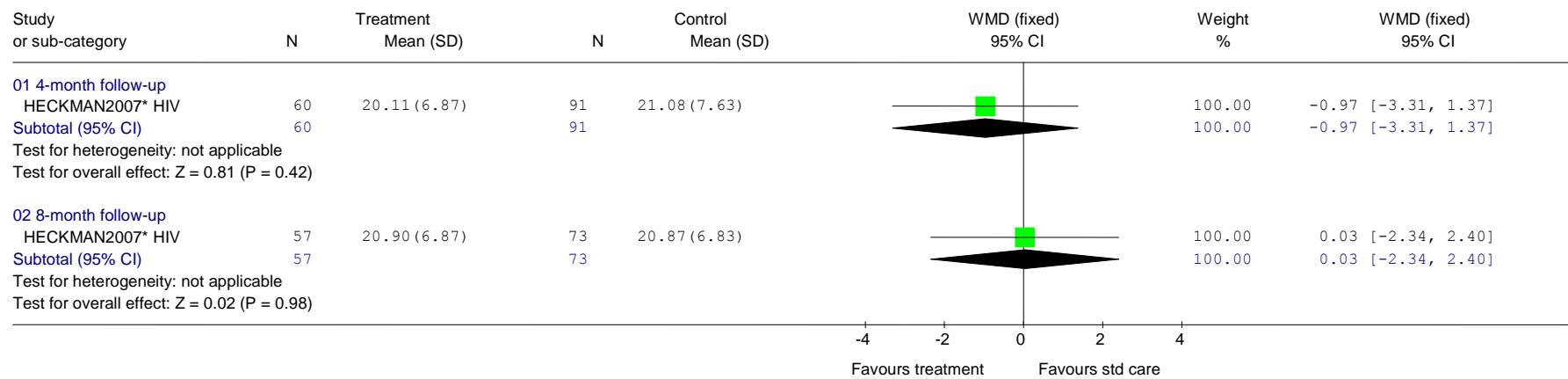


## Health education versus standard care

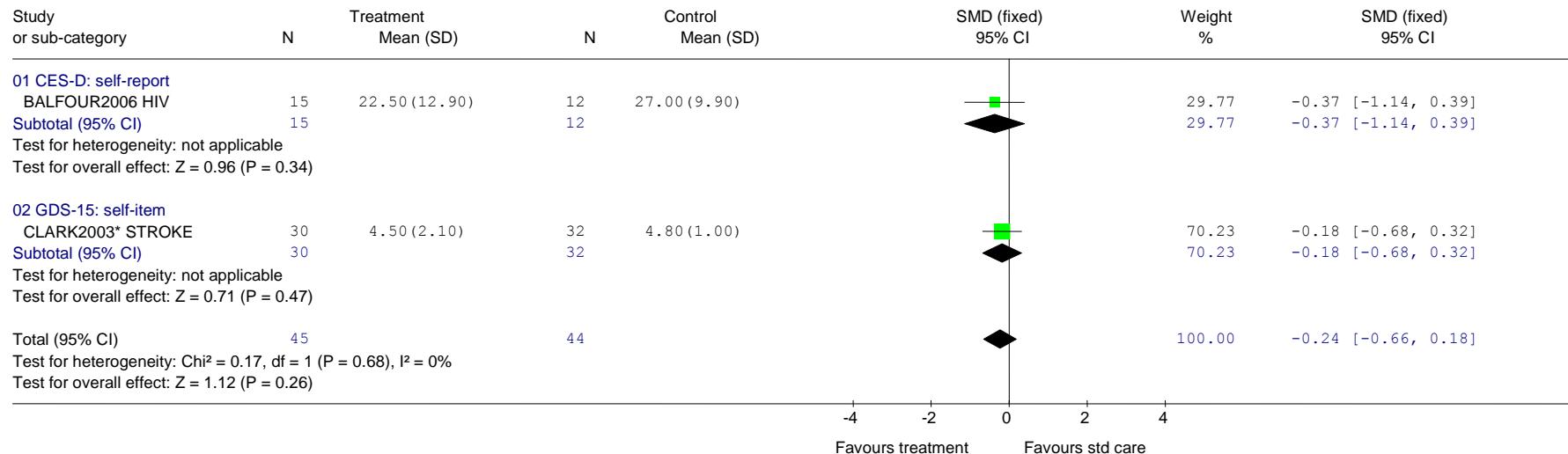
Review: DCHP: Health Education  
 Comparison: 01 Health education versus standard care  
 Outcome: 01 BDI 21 item: self-report (end of treatment)



Review: DCHP: Health Education  
 Comparison: 01 Health education versus standard care  
 Outcome: 02 BDI 21 item: self-report (follow-up)



Review: DCHP: Health Education  
 Comparison: 02 Health education plus other psychosocial component versus standard care  
 Outcome: 01 Depression (end of treatment)

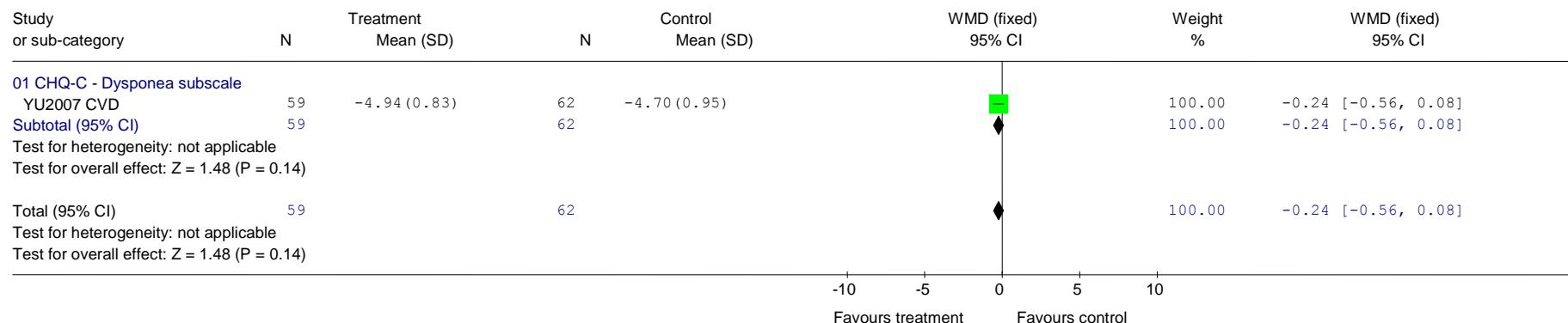


## Relaxation training versus standard care

Review: DCHP - Relaxation training  
 Comparison: 01 Relaxation versus standard care  
 Outcome: 01 Depression (Change scores)

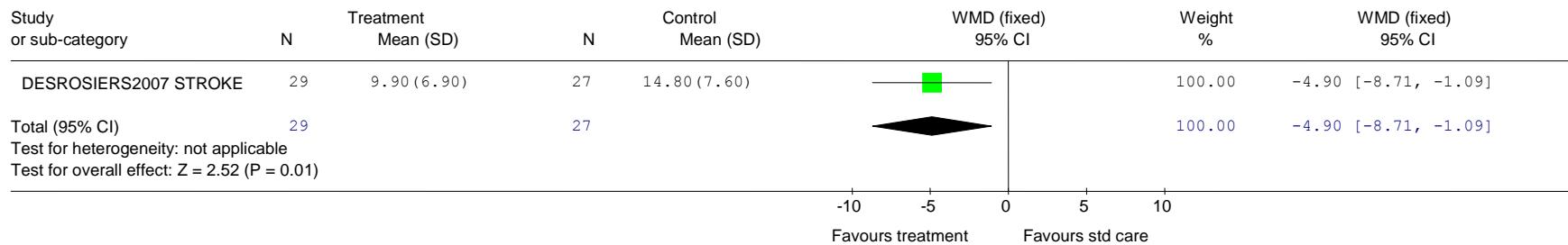


Review: DCHP - Relaxation training  
 Comparison: 01 Relaxation versus standard care  
 Outcome: 03 Health related quality of life (end of treatment)



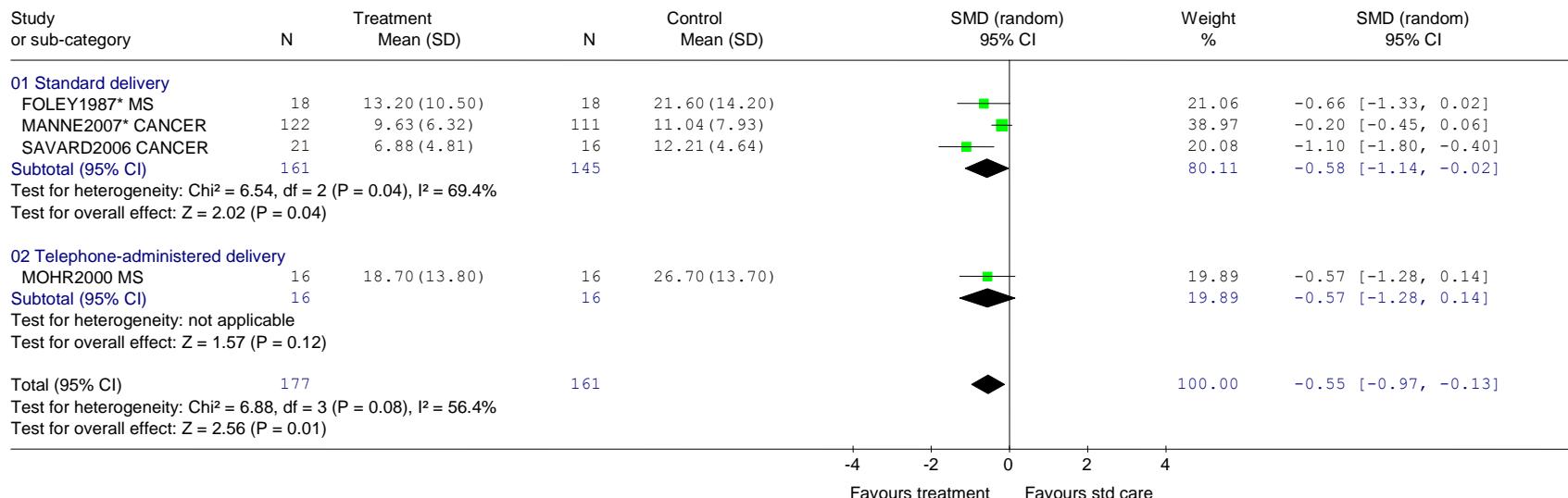
## Social support versus standard care

Review: DCHP: Social Support  
 Comparison: 01 Social Support versus standard care  
 Outcome: 01 CES-D: self-report (end of treatment)

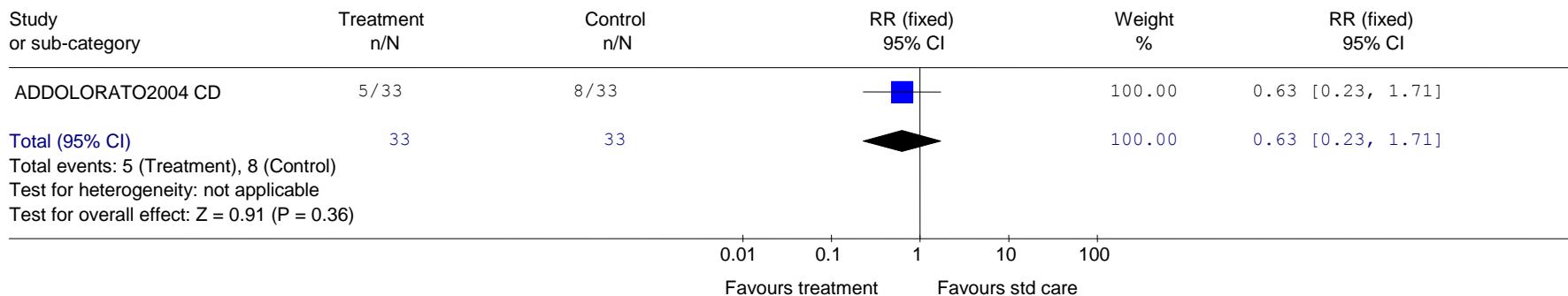


## Individual-based cognitive and behavioural interventions versus standard care

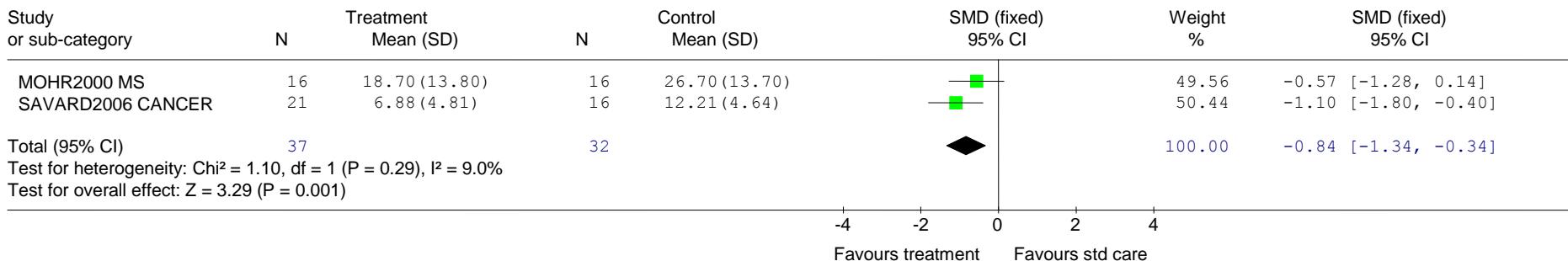
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 01 Depression (end of treatment)



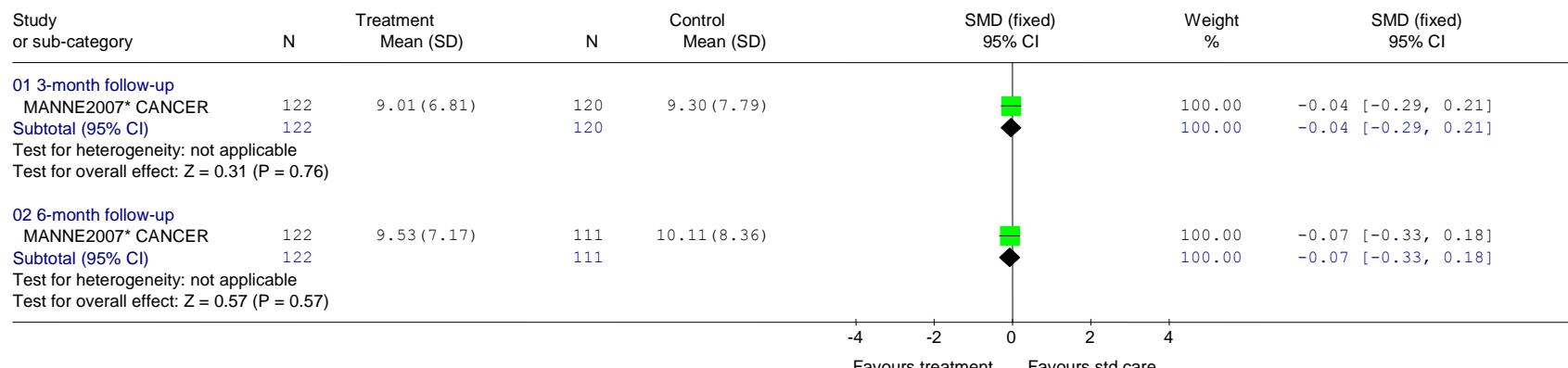
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 13 Non-remission (below cut-off)



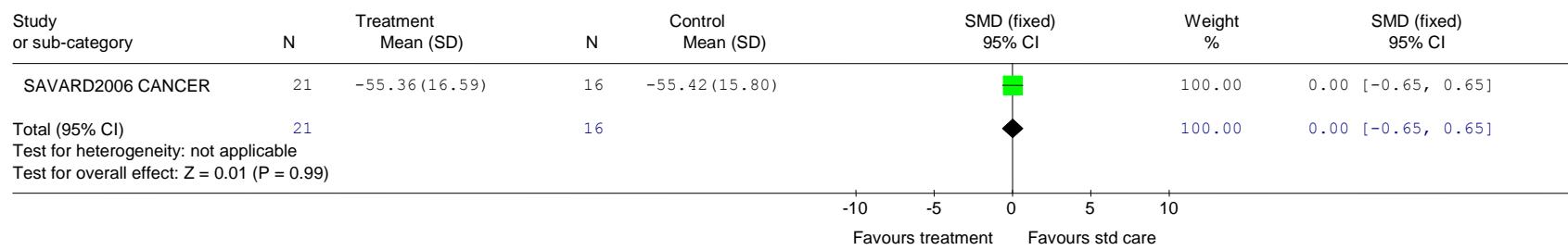
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 03 Depression (end of treatment) - sensitivity analysis - participants recruited for depression



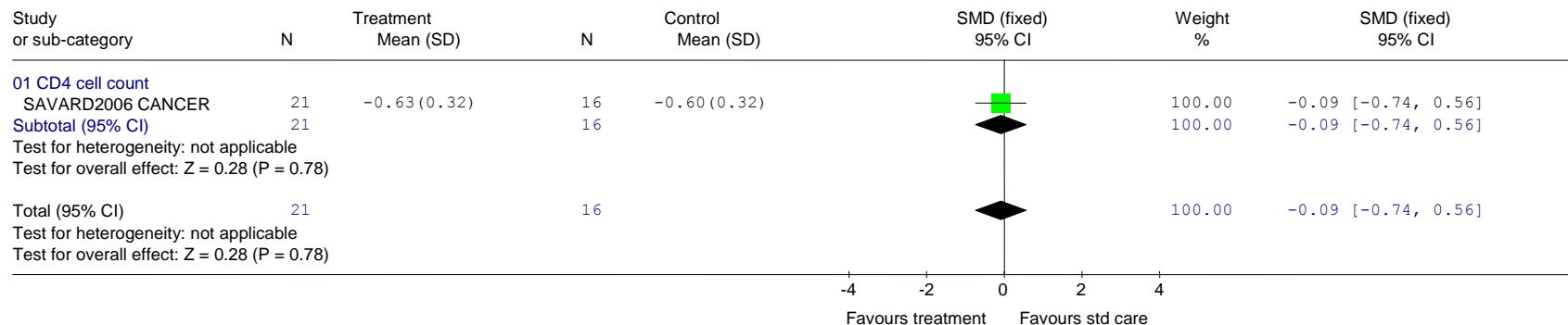
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 04 Depression (follow-up)



Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 11 QoL - QLQ-C33 (end of treatment)

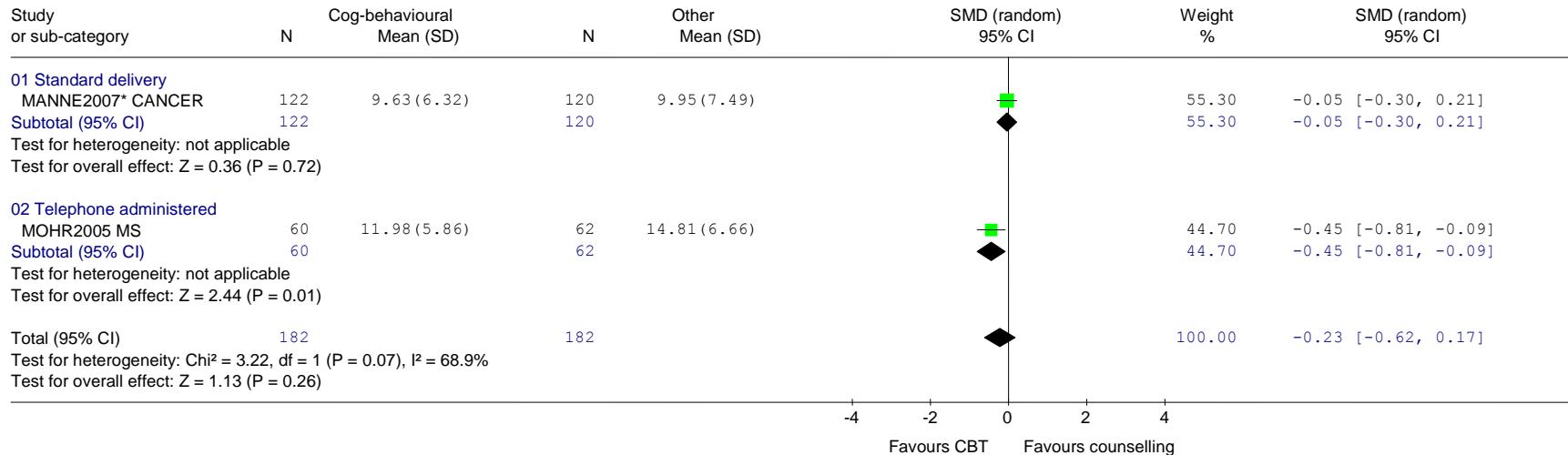


Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 01 Individual-based cognitive and behavioural intervention versus standard care  
 Outcome: 12 Physical health outcome

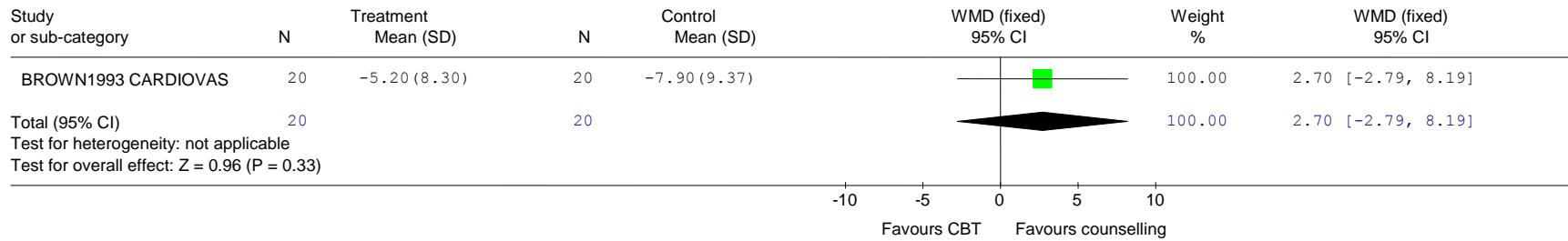


## Individual-based cognitive and behavioural interventions versus counselling

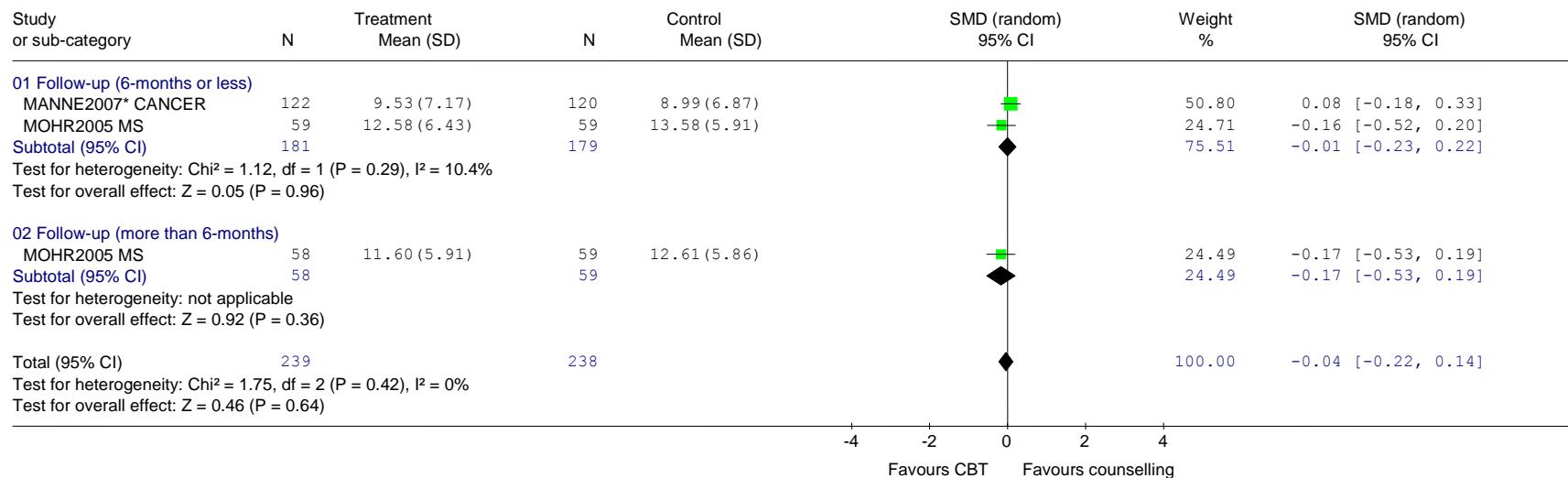
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 02 Individual-based cognitive and behavioural intervention versus counselling  
 Outcome: 01 Depression (end of treatment)



Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 02 Individual-based cognitive and behavioural intervention versus counselling  
 Outcome: 02 Depression (change score: end of treatment)



Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 02 Individual-based cognitive and behavioural intervention versus counselling  
 Outcome: 05 Depression (follow-up)

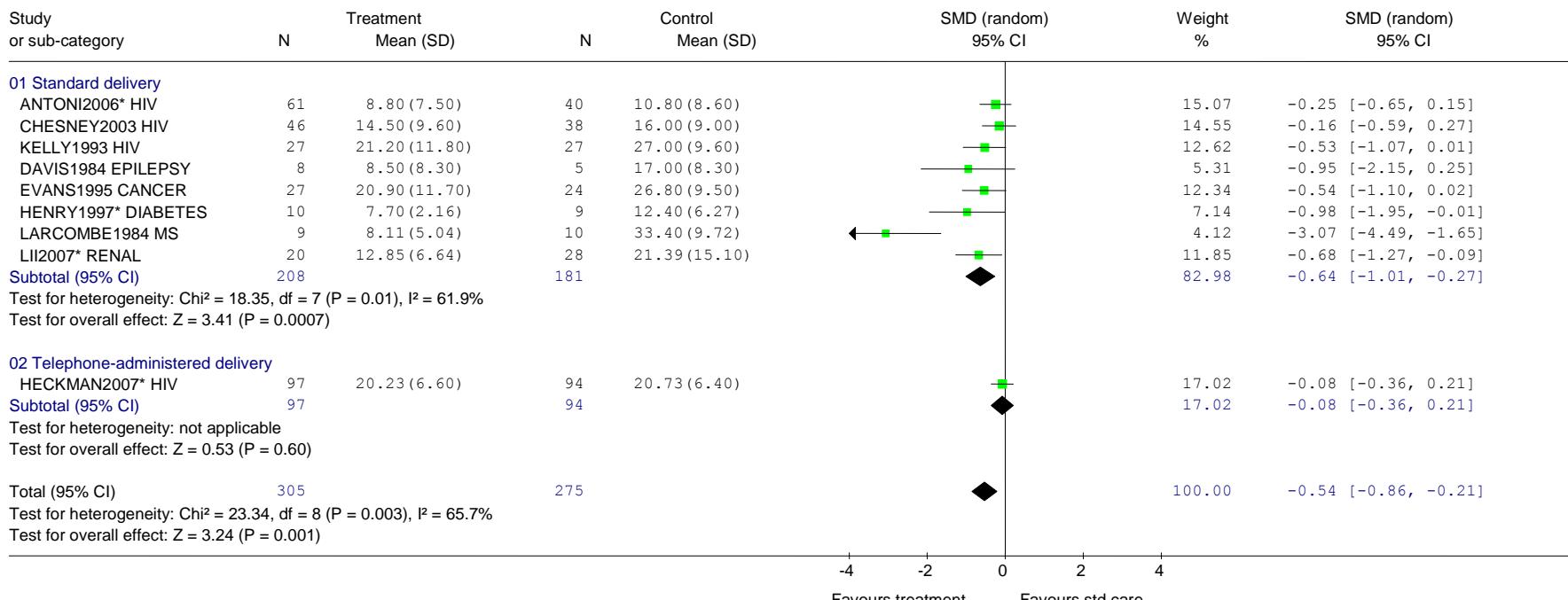


## Group-based cognitive and behavioural interventions versus standard care

Review: DCHP: Cognitive and Behavioural Interventions

Comparison: 03 Group-based cognitive and behavioural intervention versus standard care

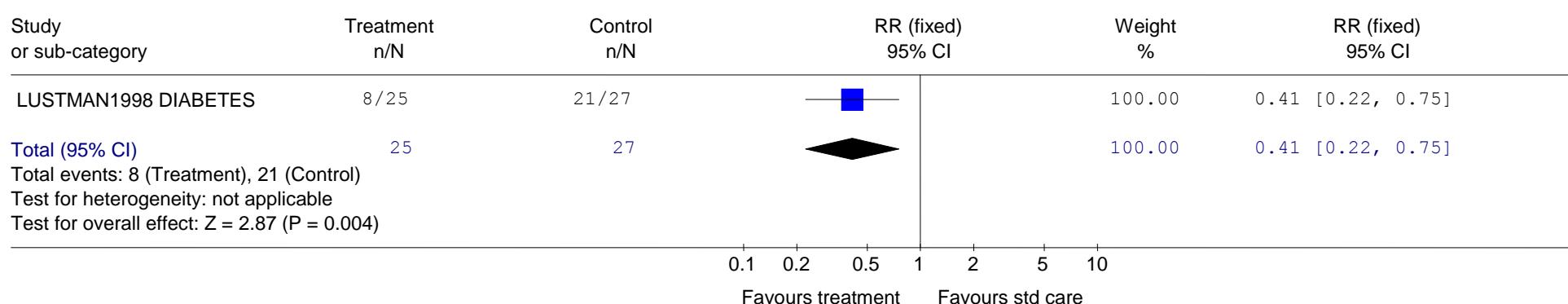
Outcome: 01 Depression (end of treatment)



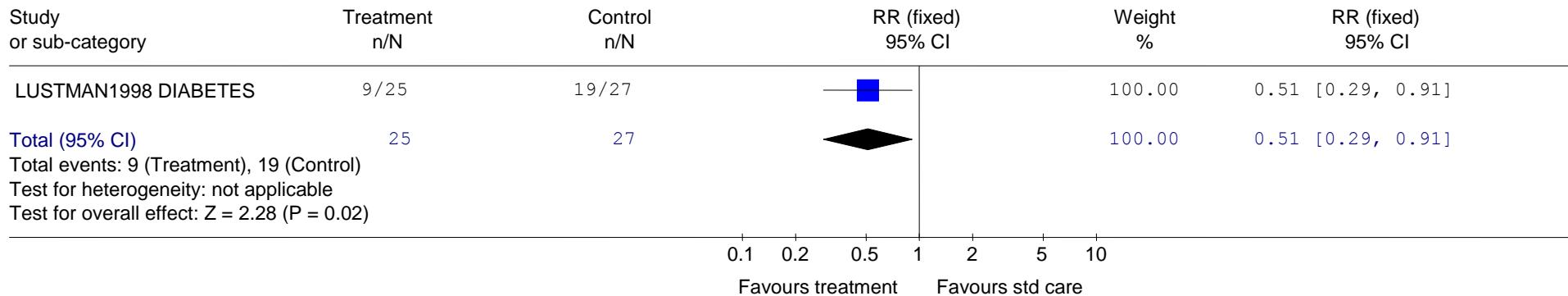
Review: DCHP: Cognitive and Behavioural Interventions

Comparison: 03 Group-based cognitive and behavioural intervention versus standard care

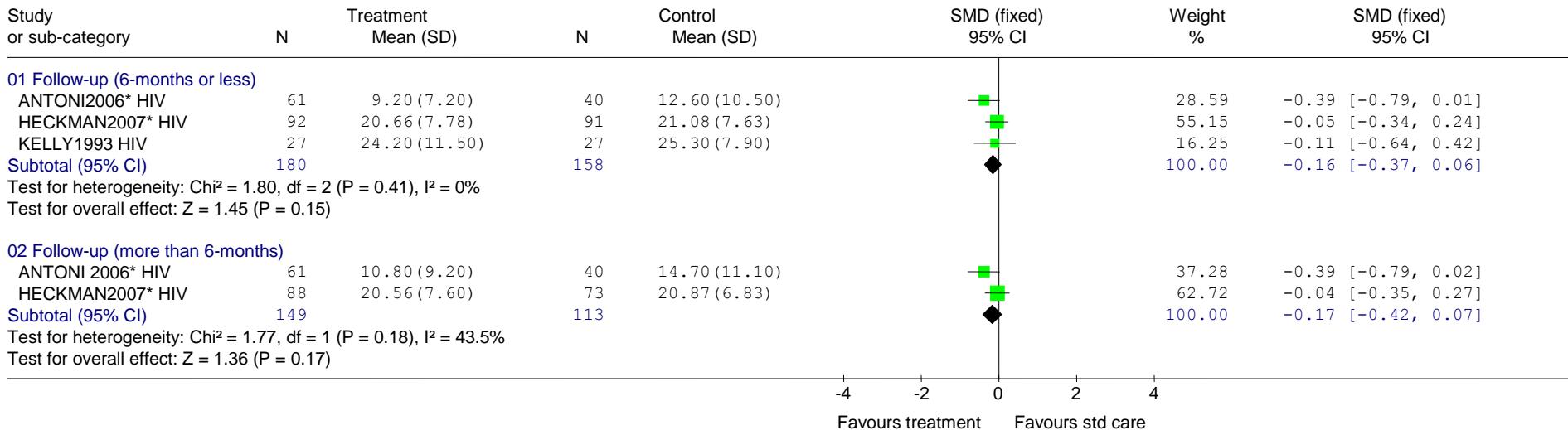
Outcome: 06 Non-remission (below cut off)



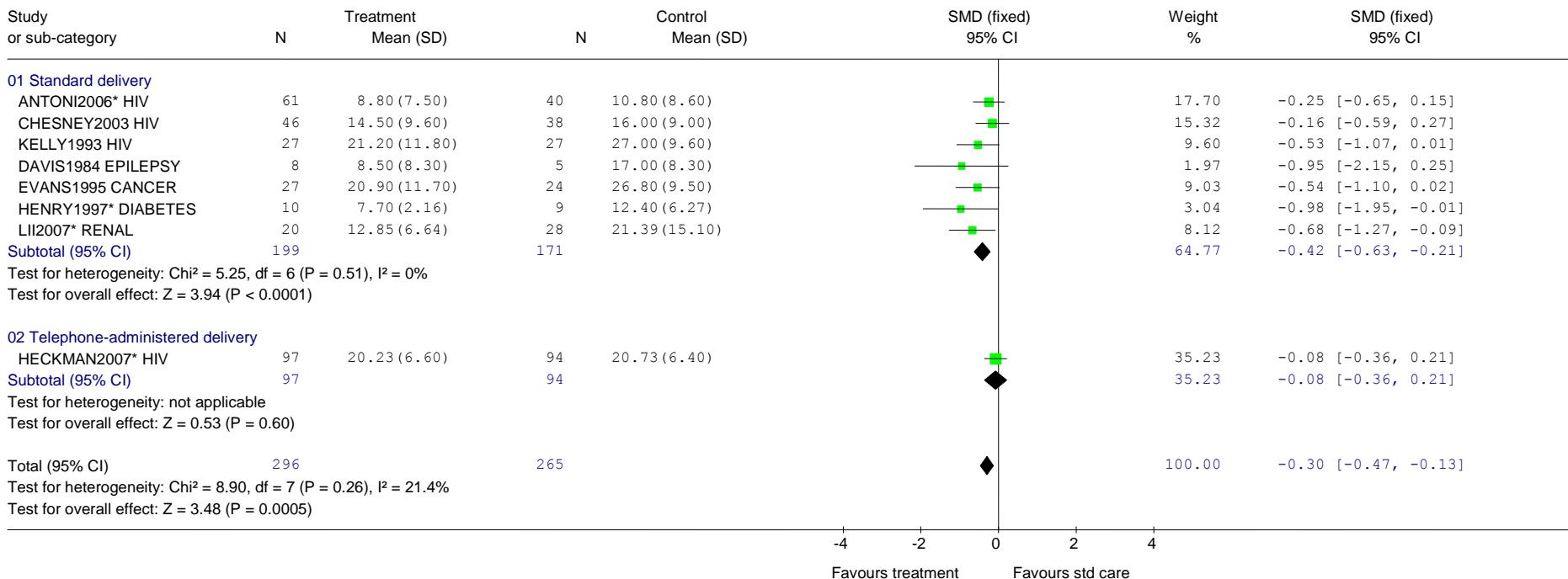
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 07 Non-response (<50% reduction from baseline)



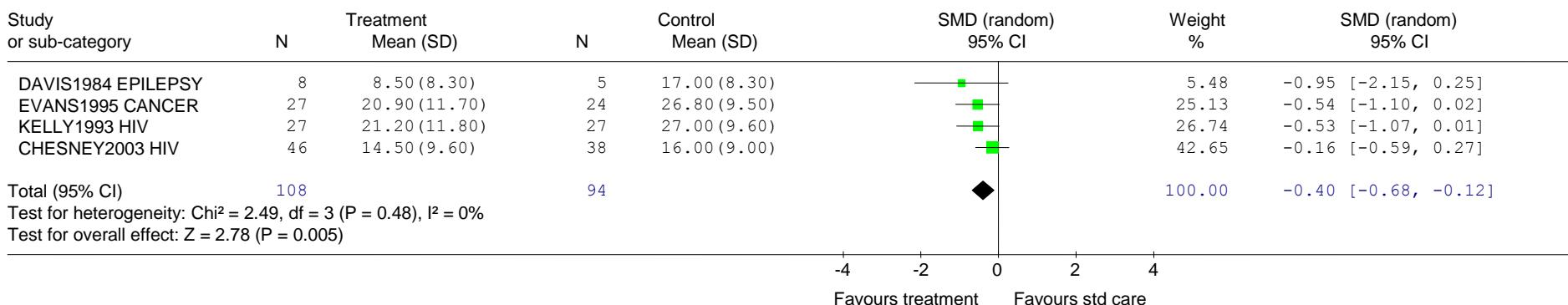
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 05 Depression (follow-up)



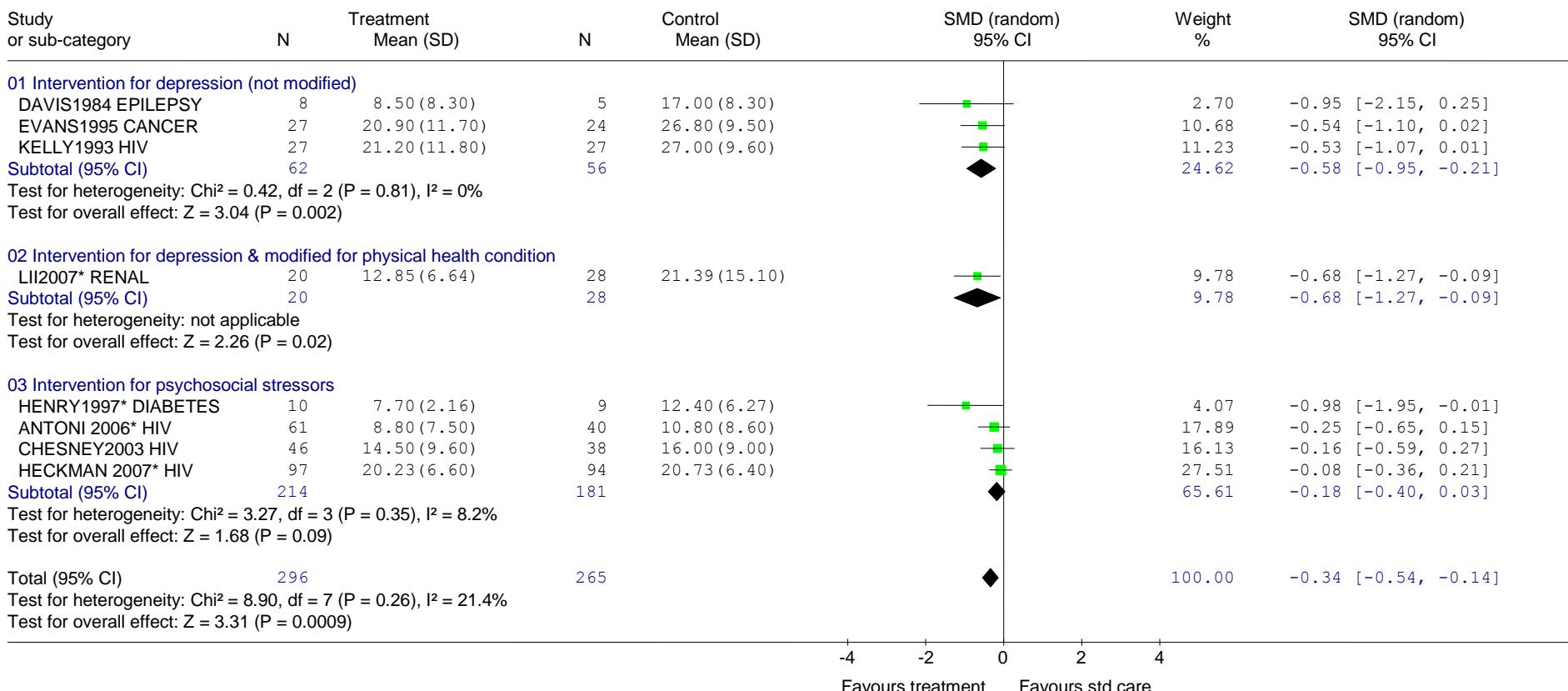
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 02 Depression (end of treatment) - sensitivity analysis (removed outlier)



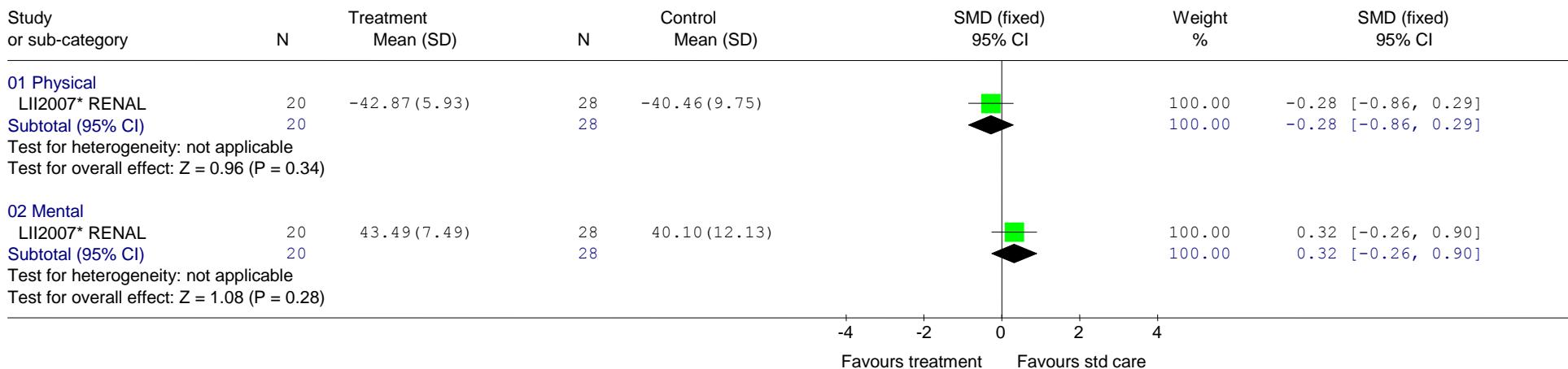
Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 04 Depression (end of treatment) - sensitivity analysis - participants recruited for depression



Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 03 Depression (end of treatment) - sub-group analysis by aim of intervention



Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 03 Group-based cognitive and behavioural intervention versus standard care  
 Outcome: 15 QoL - SF-30: self-report (end of treatment)

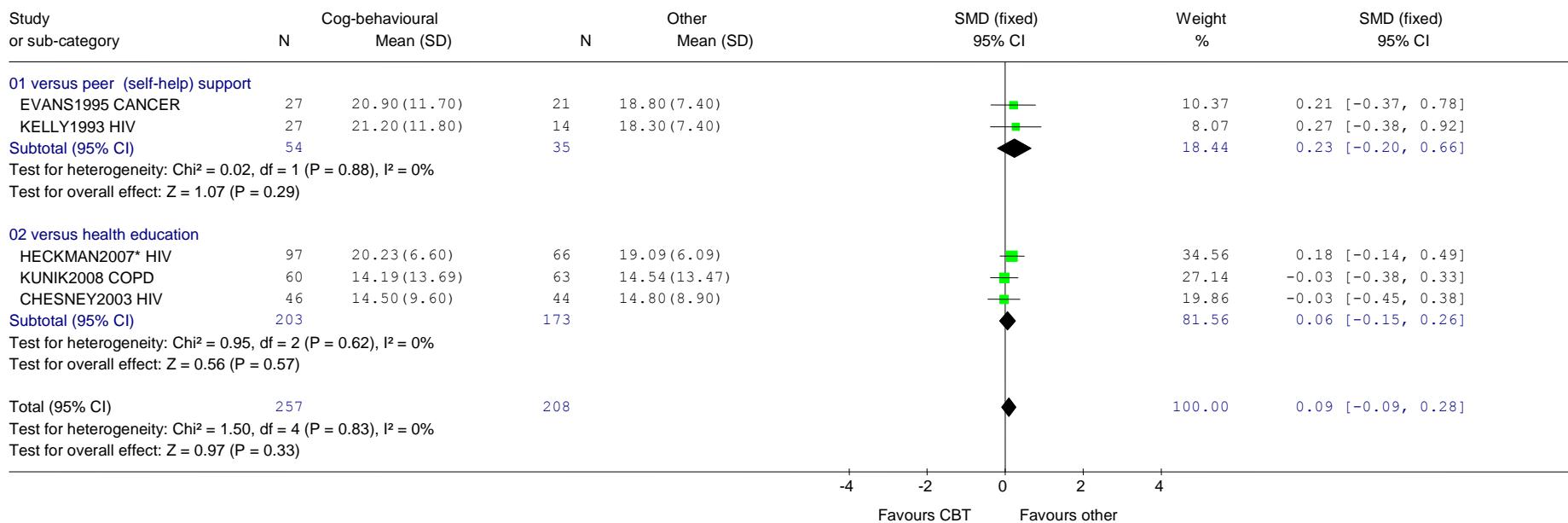


## Group-based cognitive and behavioural interventions versus other (peer self-help support and health education)

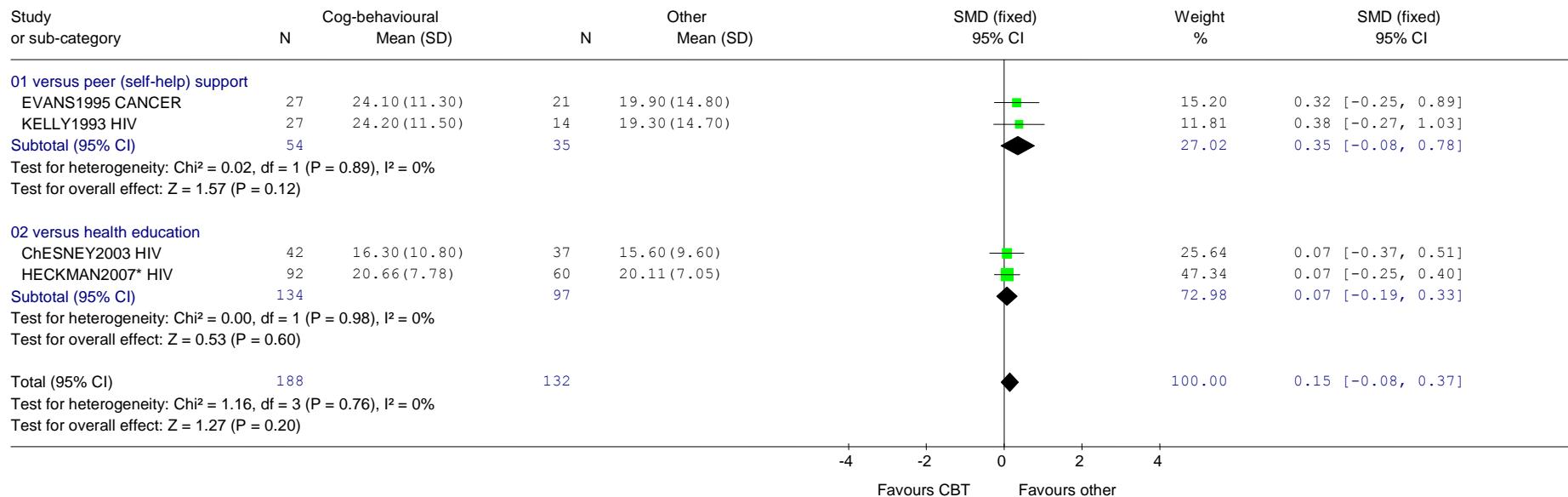
Review: DCHP: Cognitive and Behavioural Interventions

Comparison: 06 Group-based cognitive and behavioural intervention versus other (peer self-help support and health education)

Outcome: 01 Depression (end of treatment)

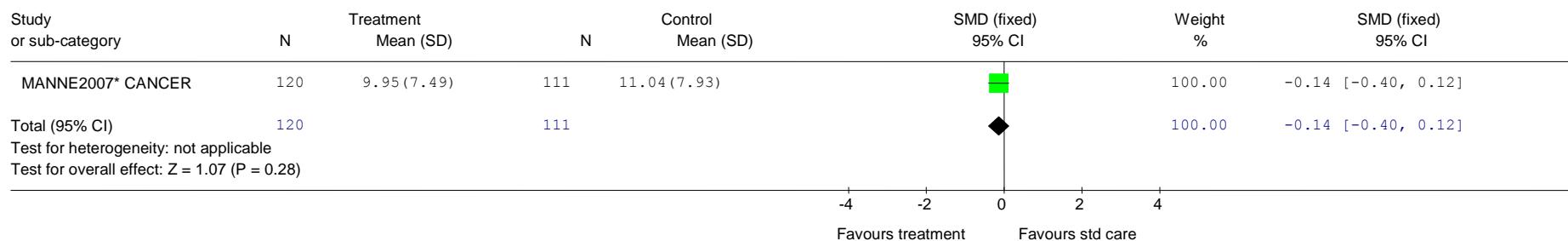


Review: DCHP: Cognitive and Behavioural Interventions  
 Comparison: 06 Group-based cognitive and behavioural intervention versus other (peer self-help support and health education)  
 Outcome: 02 Depression at follow-up (6-months of less)



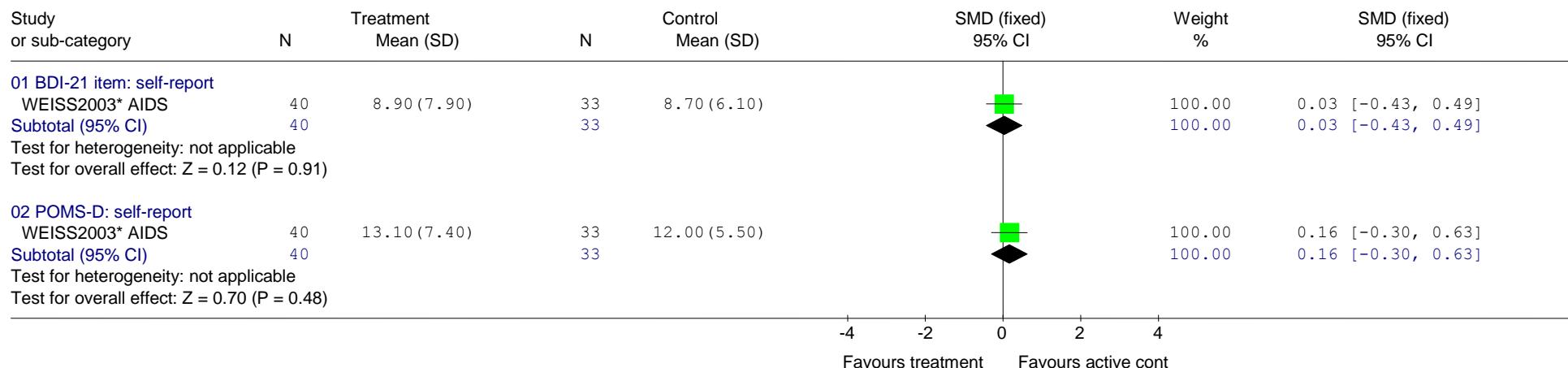
## Counselling versus standard care

Review: DCHP: Counselling  
 Comparison: 01 Counselling versus standard care  
 Outcome: 01 BDI: self report (end of treatment)

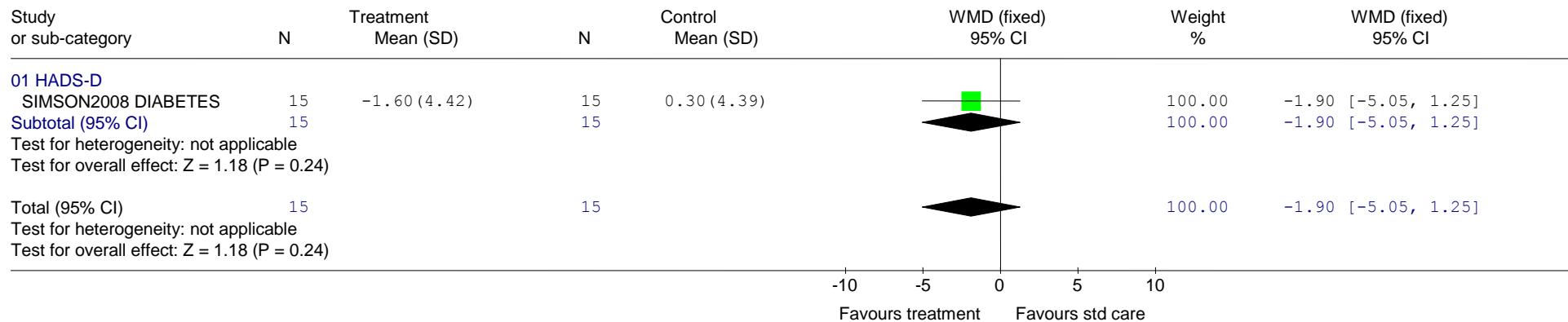


## Group existential therapy versus active control

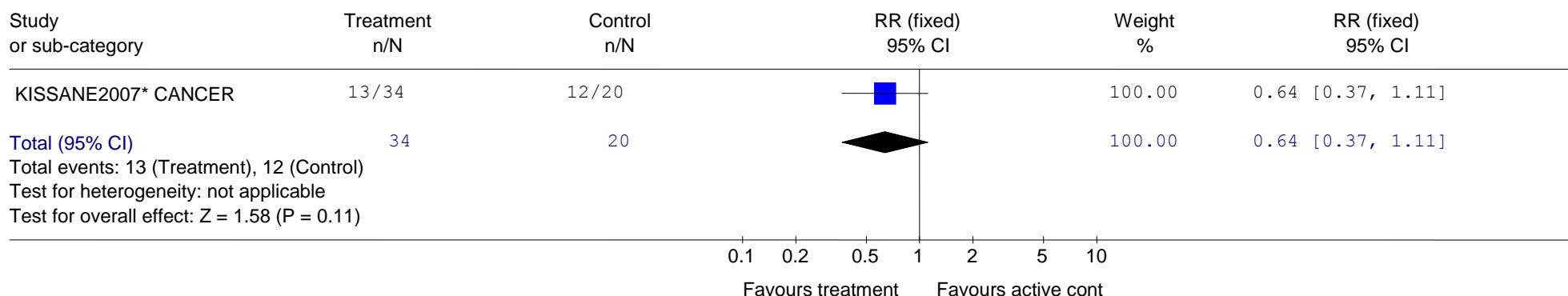
Review: DCHP: Group Existential Therapy  
 Comparison: 01 Group existential therapy  
 Outcome: 01 Depression (end of treatment)



Review: DCHP: Group Existential Therapy  
 Comparison: 01 Group existential therapy  
 Outcome: 03 Depression (Change score at end of treatment)



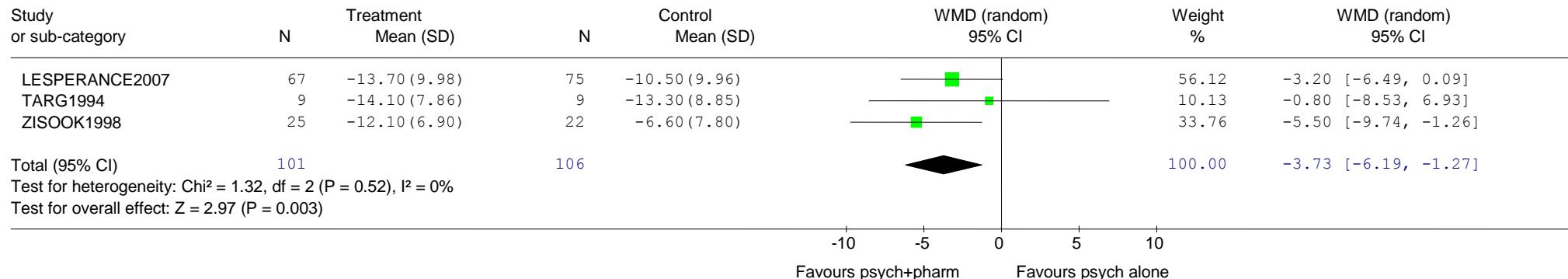
Review: DCHP: Group Existential Therapy  
 Comparison: 01 Group existential therapy  
 Outcome: 05 Non-remission (still meeting diagnosis of depression) - end of treatment



## Combination interventions

### Psychological/psychosocial (psych) interventions plus pharmacological interventions (pharm) versus psychological/psychosocial intervention alone

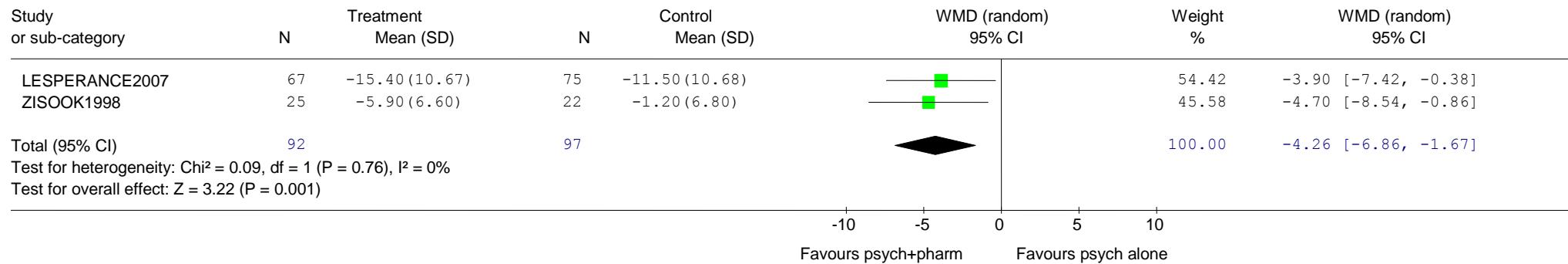
Review: DCHP: Psych + Pharm  
 Comparison: 01 SSRIs + psych versus psych  
 Outcome: 01 Depression (HAM-D)



Test for heterogeneity:  $\text{Chi}^2 = 1.32$ , df = 2 ( $P = 0.52$ ),  $I^2 = 0\%$   
 Test for overall effect:  $Z = 2.97$  ( $P = 0.003$ )

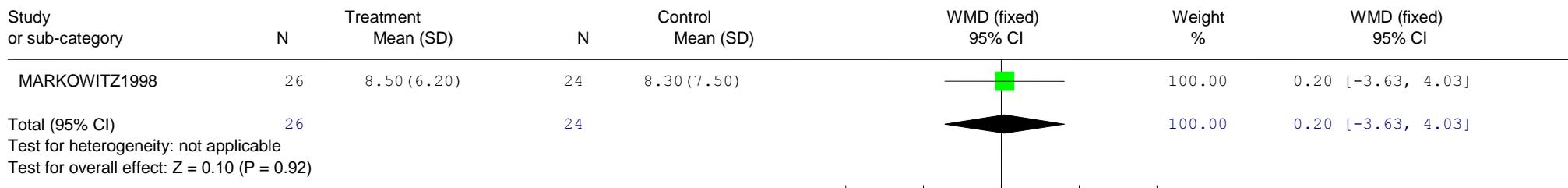
Review: DCHP: Psych + Pharm  
 Comparison: 01 SSRIs + psych versus psych

Outcome: 02 Depression (BDI)

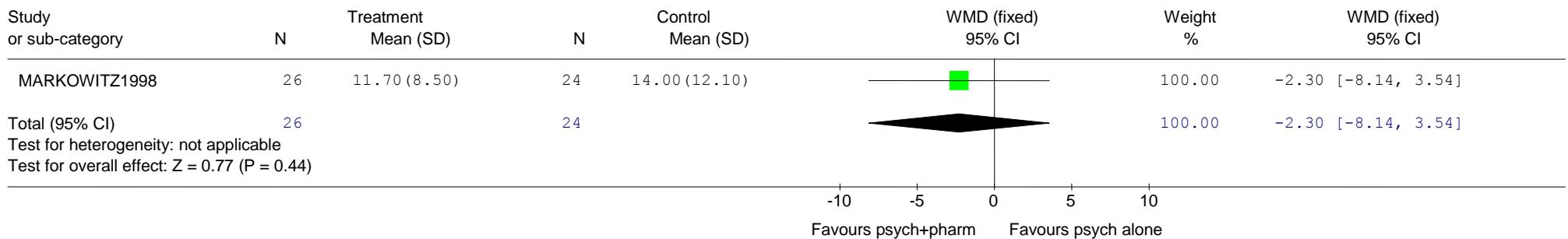


Test for heterogeneity:  $\text{Chi}^2 = 0.09$ , df = 1 ( $P = 0.76$ ),  $I^2 = 0\%$   
 Test for overall effect:  $Z = 3.22$  ( $P = 0.001$ )

Review: DCHP: Psych + Pharm  
 Comparison: 02 TCAs + psych versus psychl  
 Outcome: 01 Depression (HAM-D)

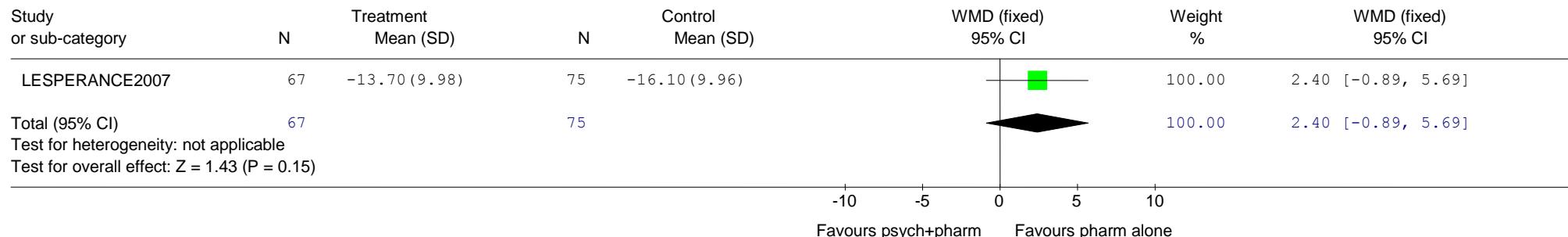


Review: DCHP: Psych + Pharm  
 Comparison: 02 TCAs + psych versus psych  
 Outcome: 02 Depression (BDI)

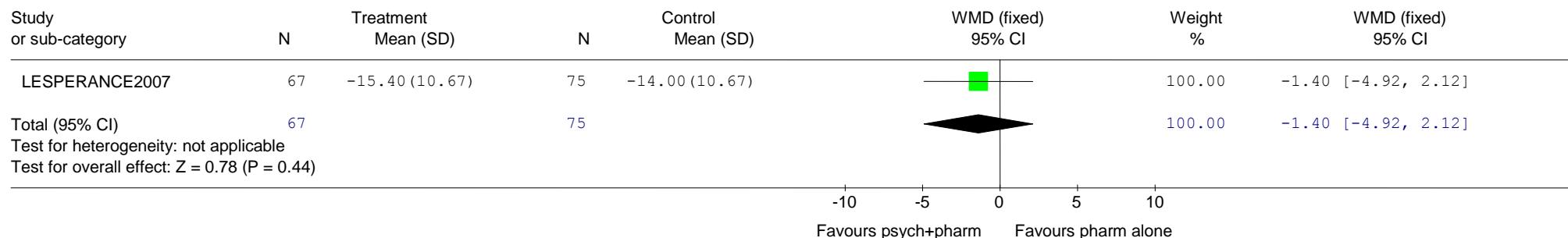


## Psychological/psychosocial interventions (psych) plus pharmacological interventions (pharm) versus pharmacological interventions alone

Review: DCHP: Psych + Pharm  
 Comparison: 03 SSRI + Psych versus SSRI  
 Outcome: 01 Depression (HAM-D)



Review: DCHP: Psych + Pharm  
 Comparison: 03 SSRI + Psych versus SSRI  
 Outcome: 02 Depression (BDI)



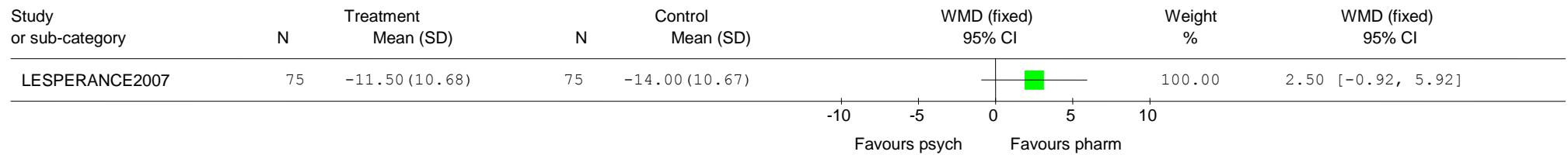
## *Psychological/psychosocial versus pharmacological interventions*

### **Psychological/psychosocial interventions (psych) versus pharmacological interventions (pharm)**

Review: DCHP: Psych + Pharm  
 Comparison: 04 Psych versus SSRIs  
 Outcome: 01 Depression (HAM-D)

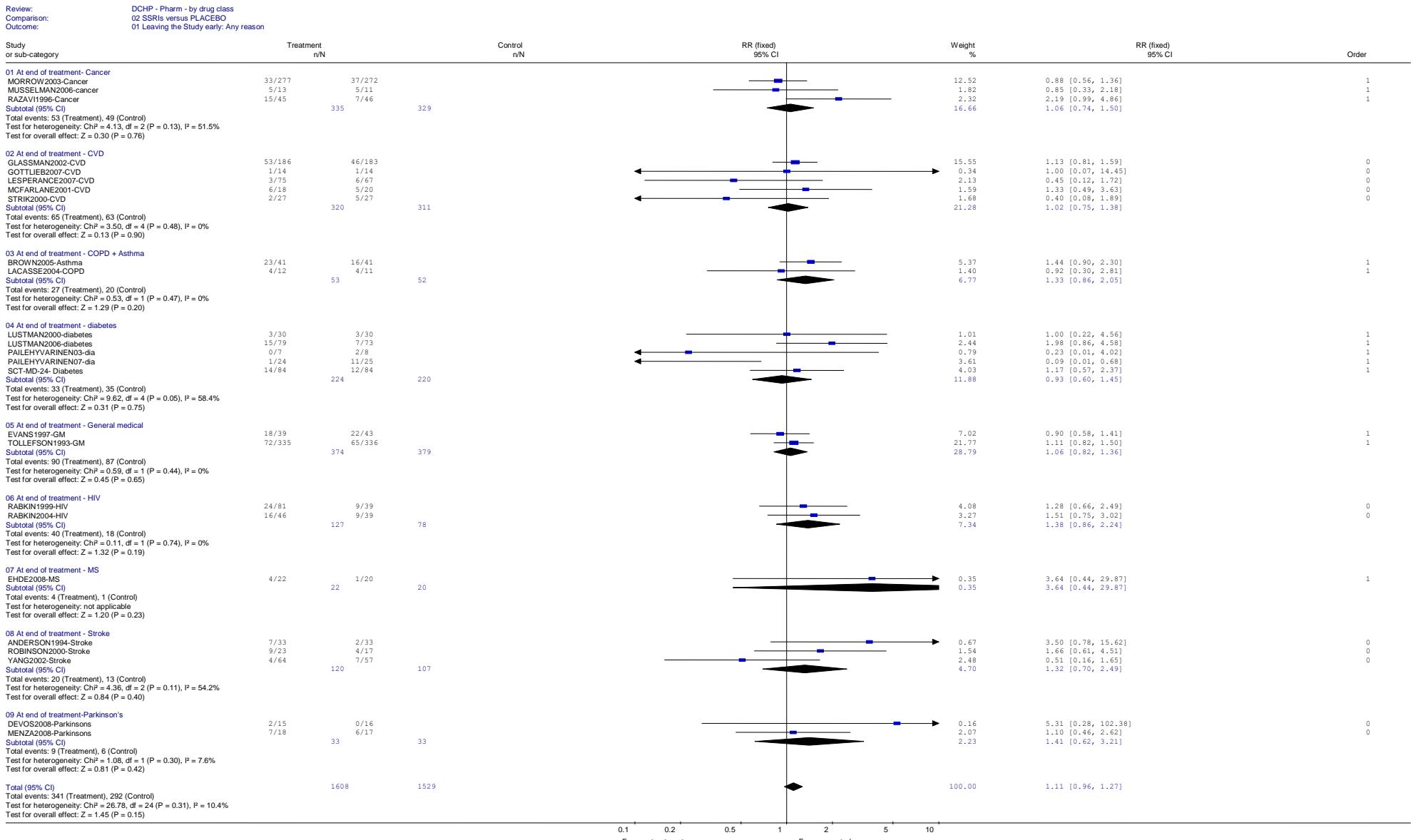


Review: DCHP: Psych + Pharm  
 Comparison: 04 Psych versus SSRIs  
 Outcome: 02 Depression (BDI)

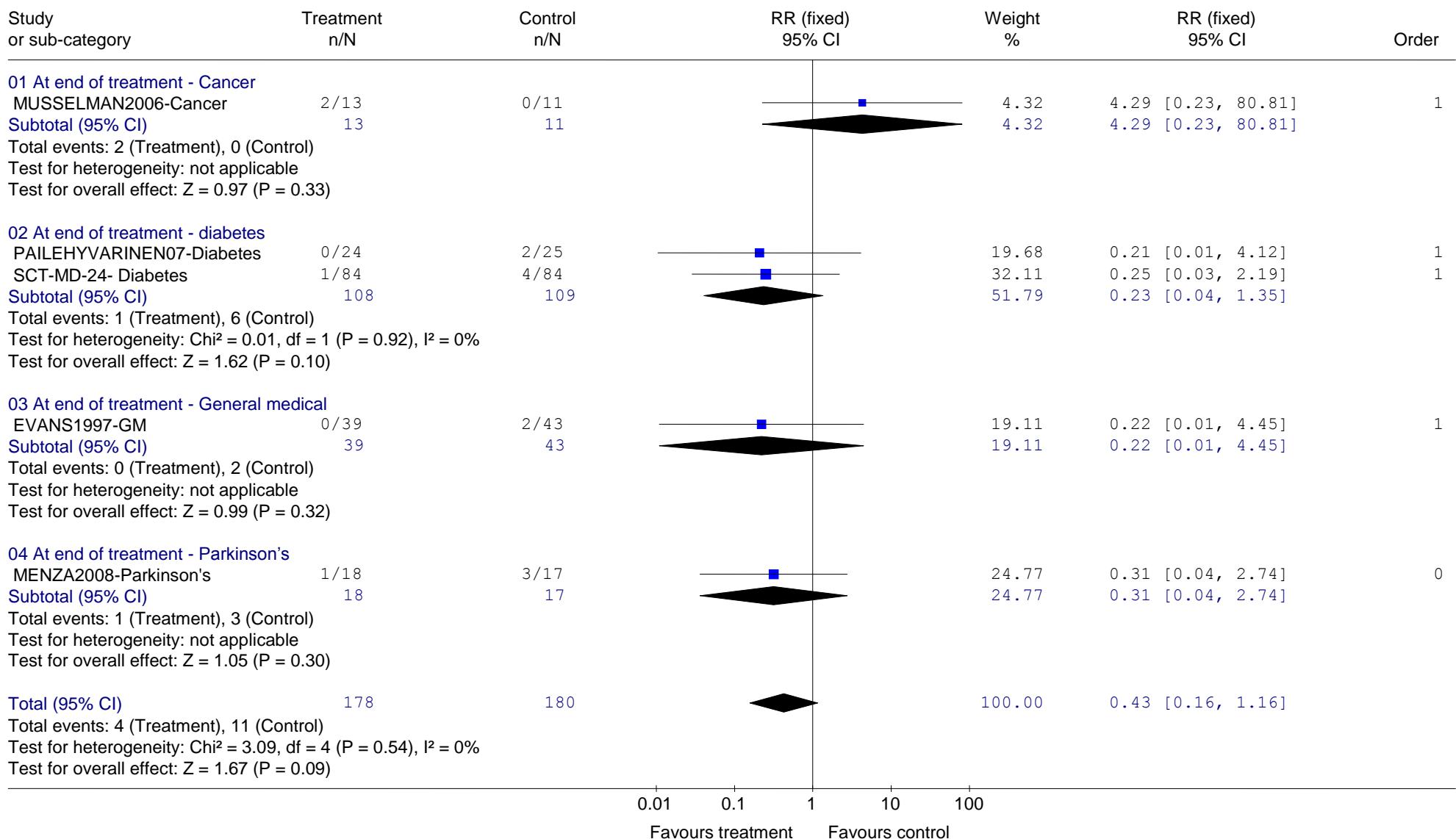


# Pharmacological interventions

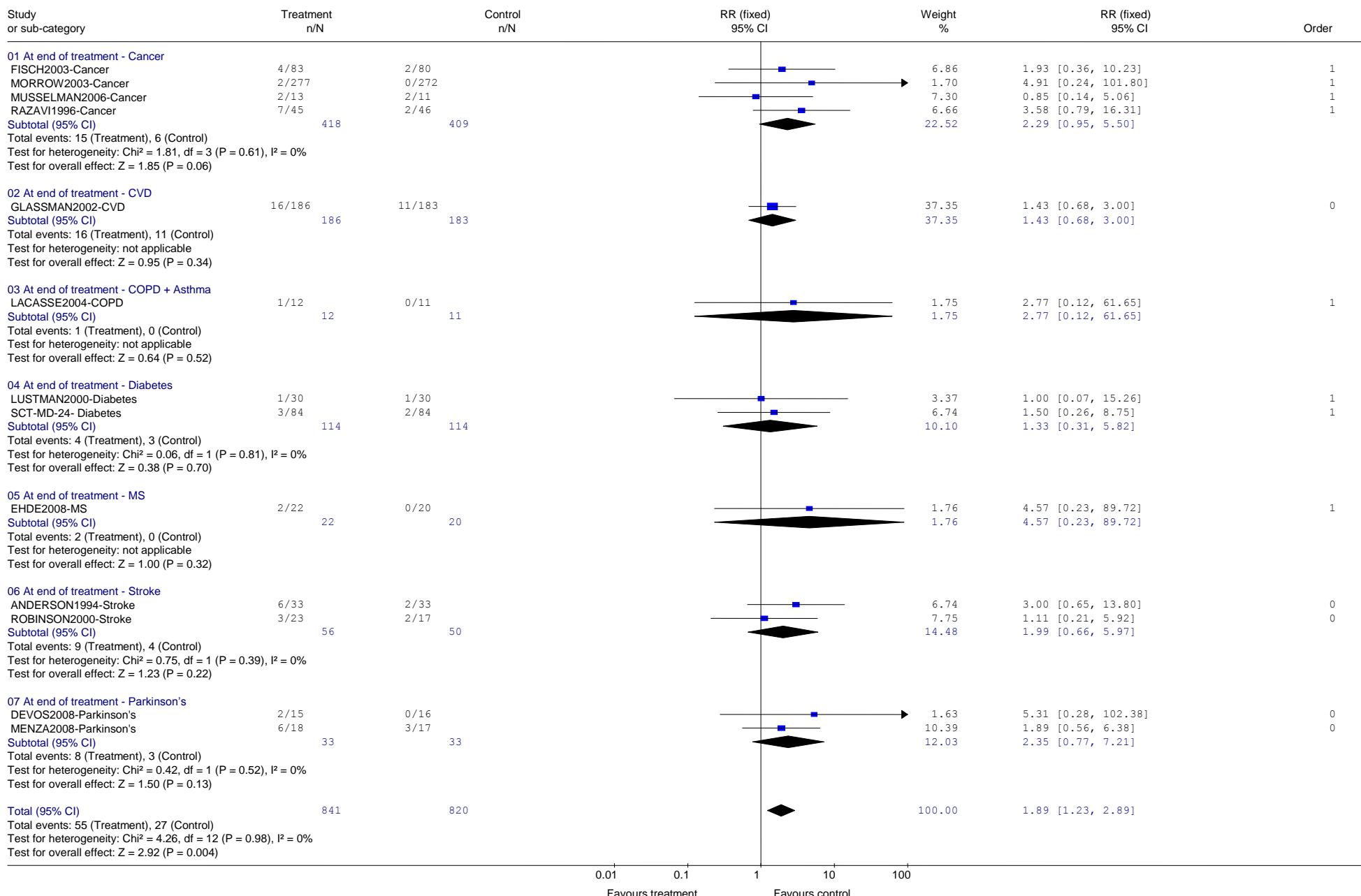
## SSRIs versus placebo



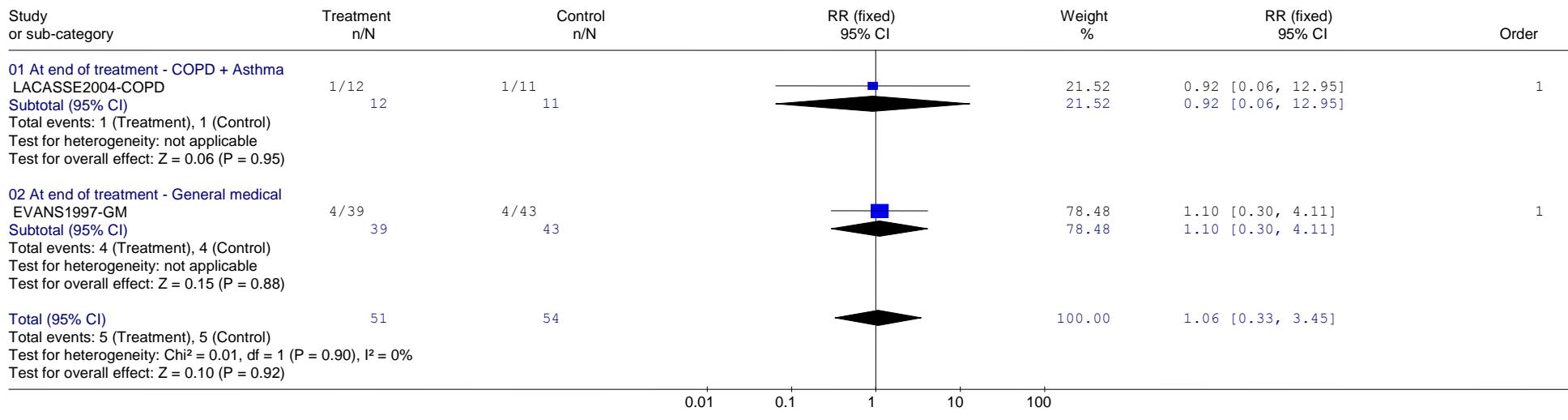
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 02 Leaving the study early: Lack of efficacy



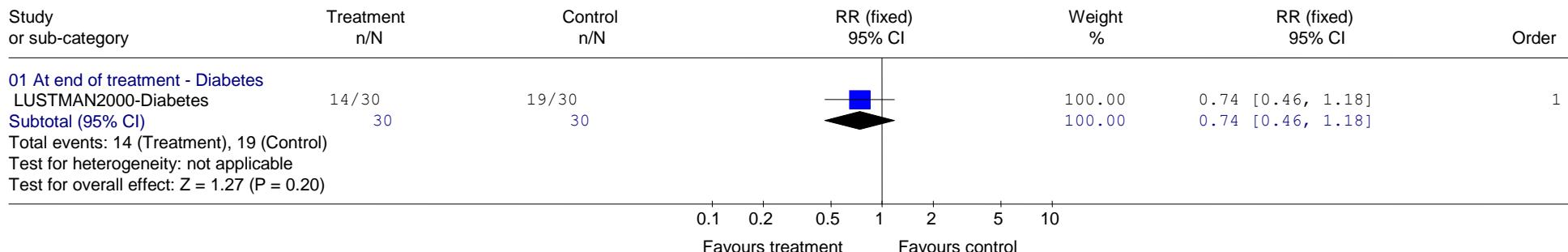
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 03 Leaving the Study early: Due to adverse events



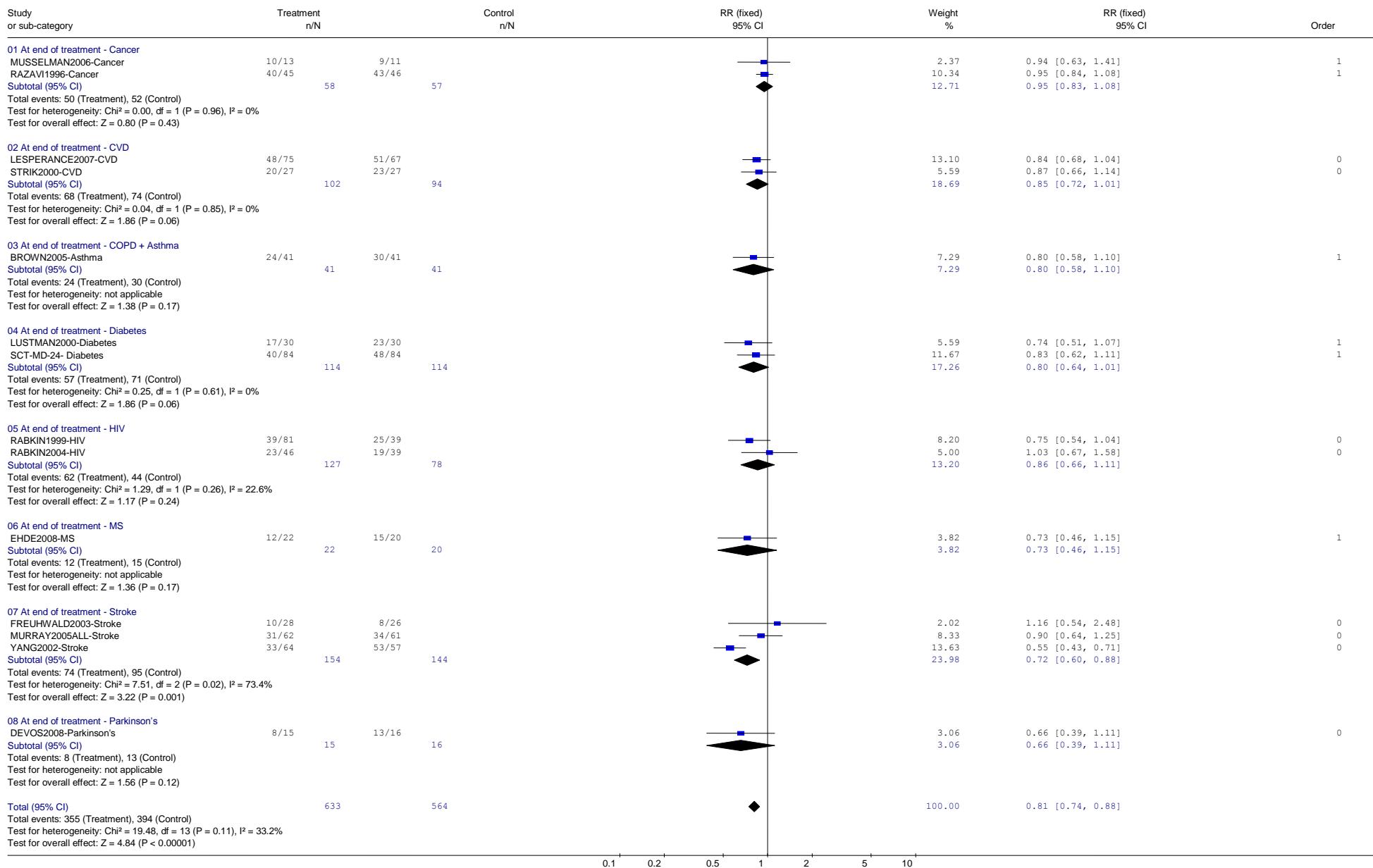
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 10 Mortality



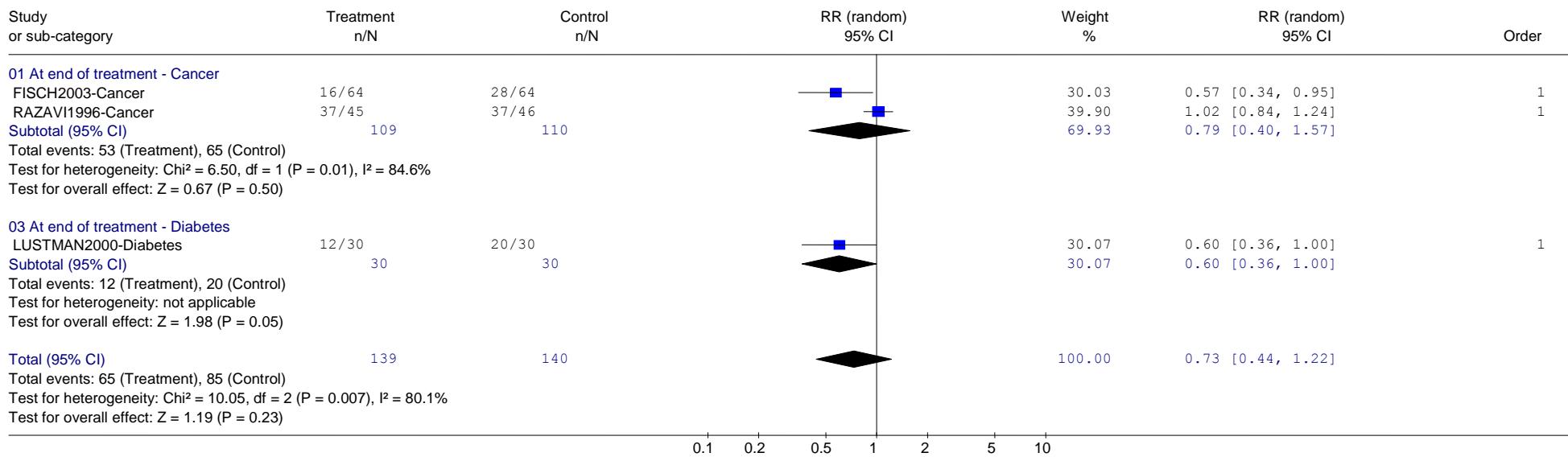
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 30 Depression: 1. Not achieving success/ remission (reaching a specified cut-off) - patient rated



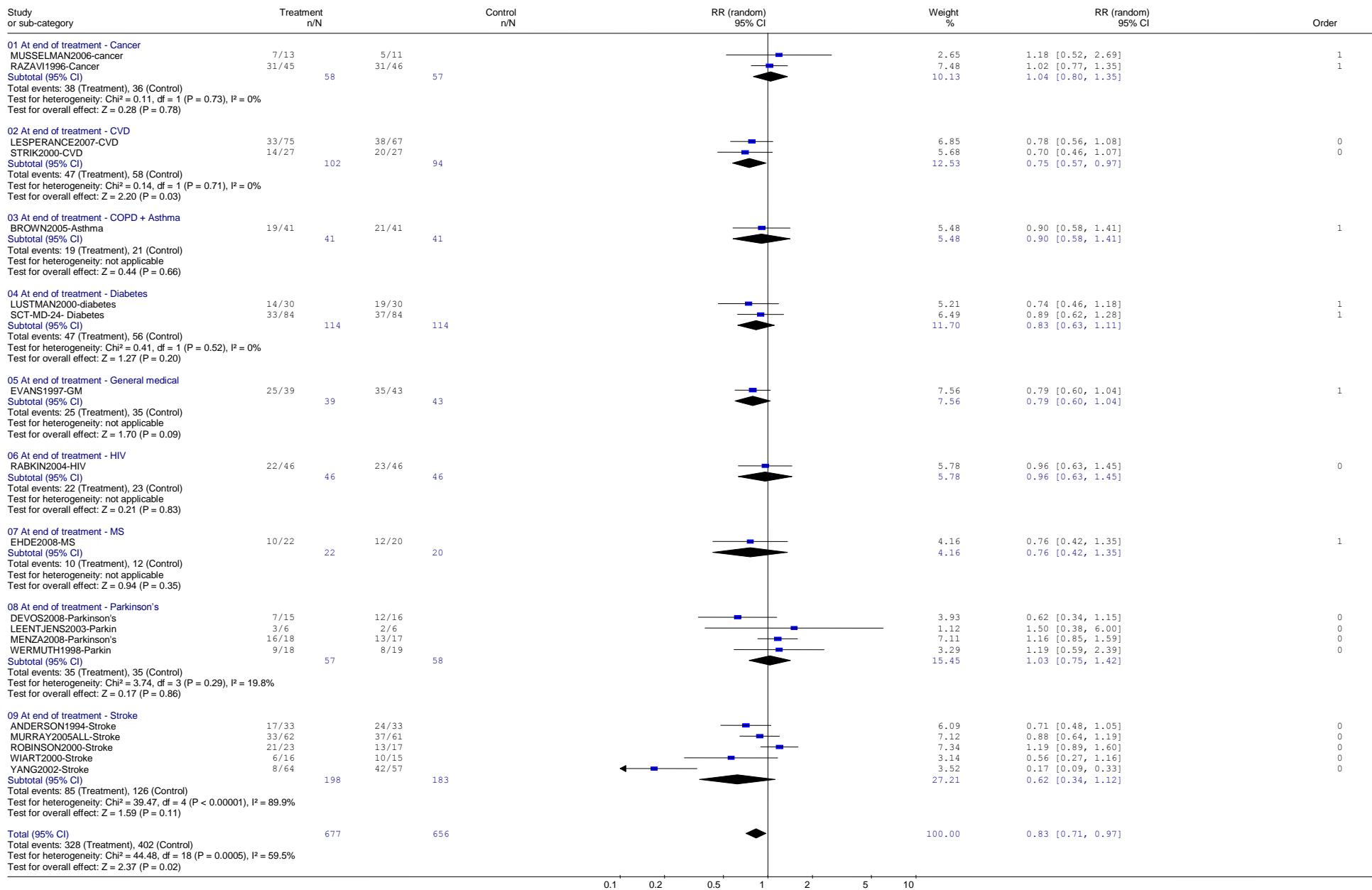
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 31 Depression: 1. Not achieving success/ remission (reaching a specified cut-off) - observer rated



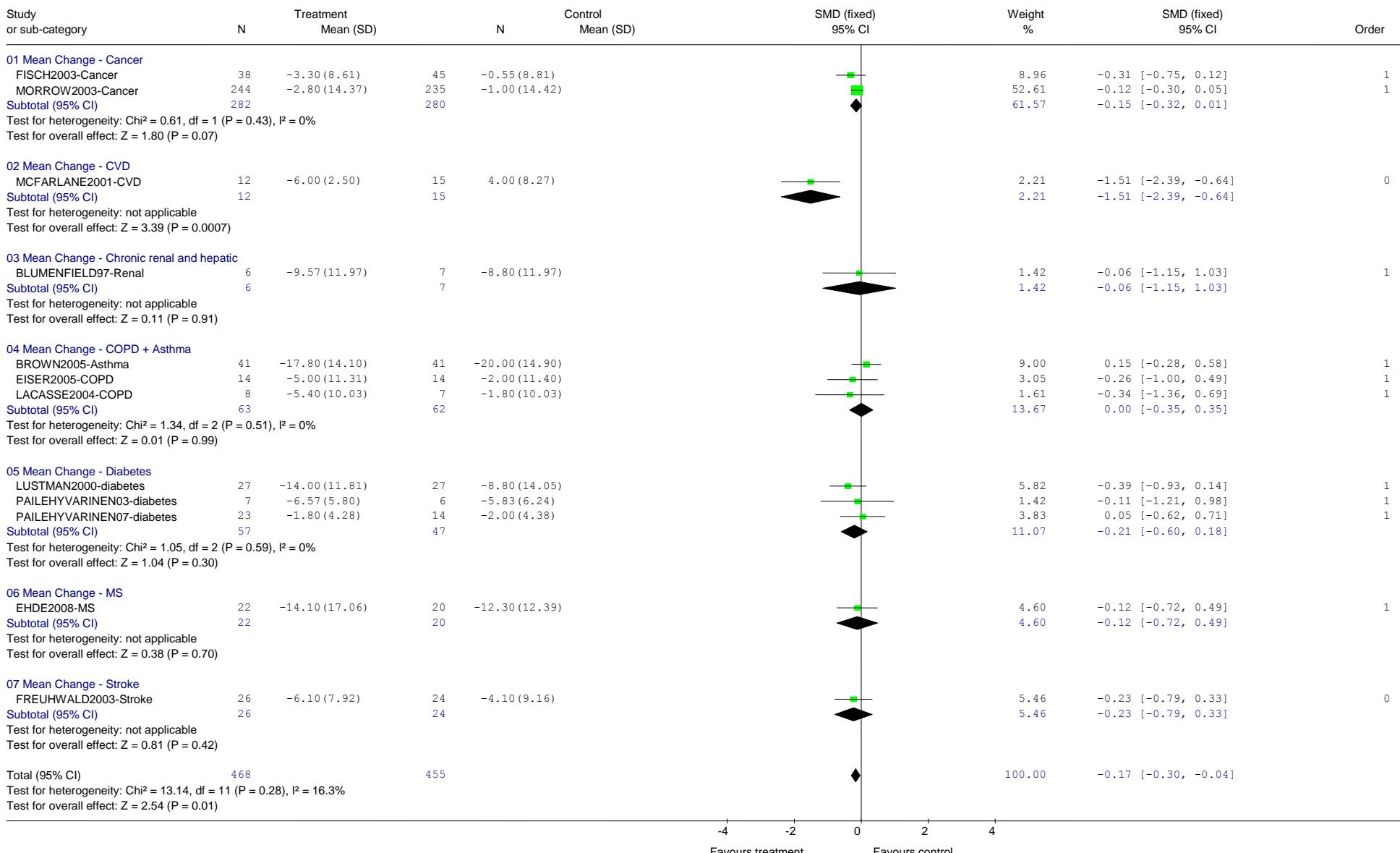
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 32 Depression: 2. Non-response (not achieving 50% reduction from baseline) - patient rated e.g. BDI



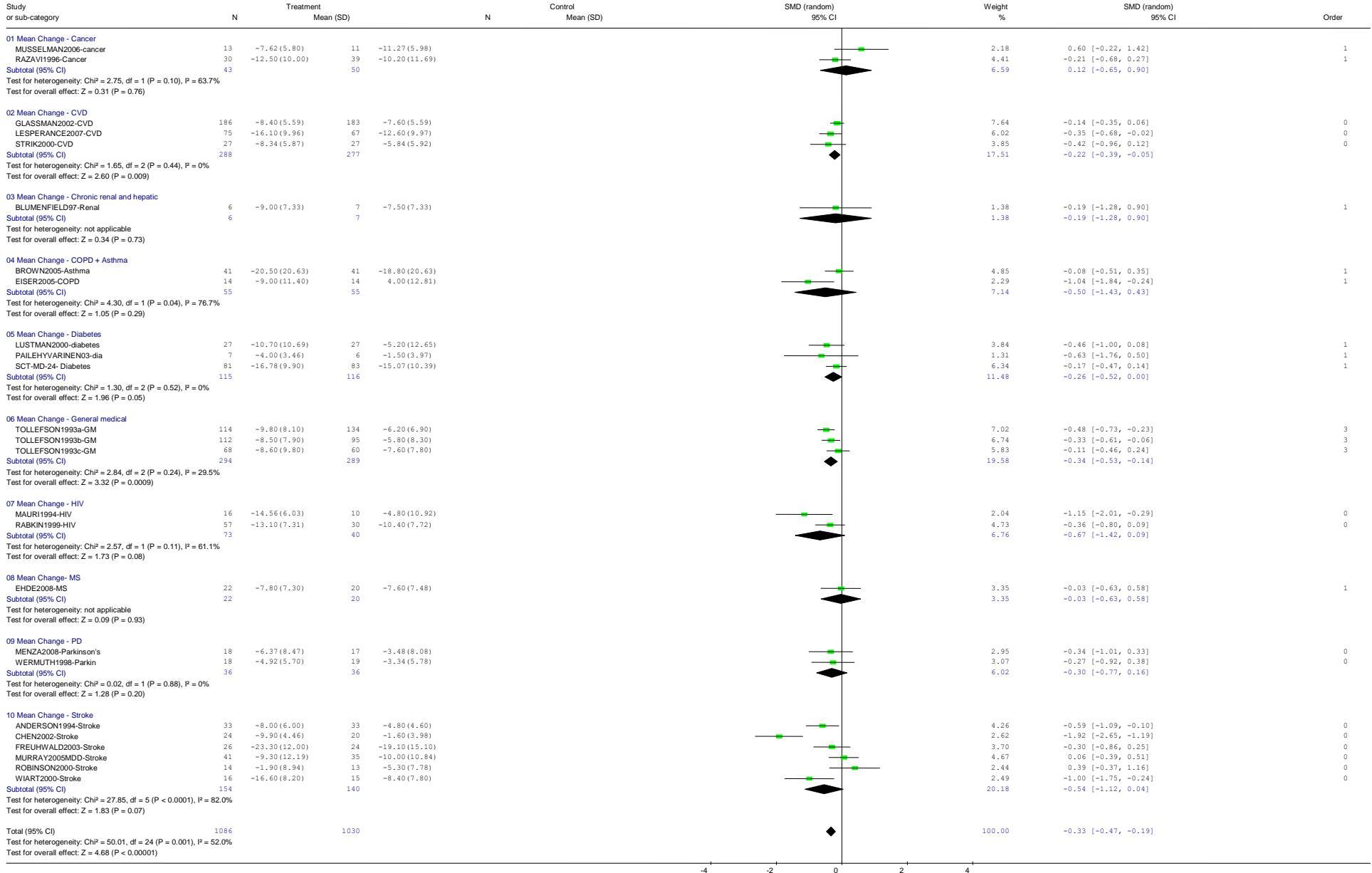
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 33 Depression: 2. Non-response (not achieving 50% reduction from baseline) - observer rated e.g. HAMD, MADRS



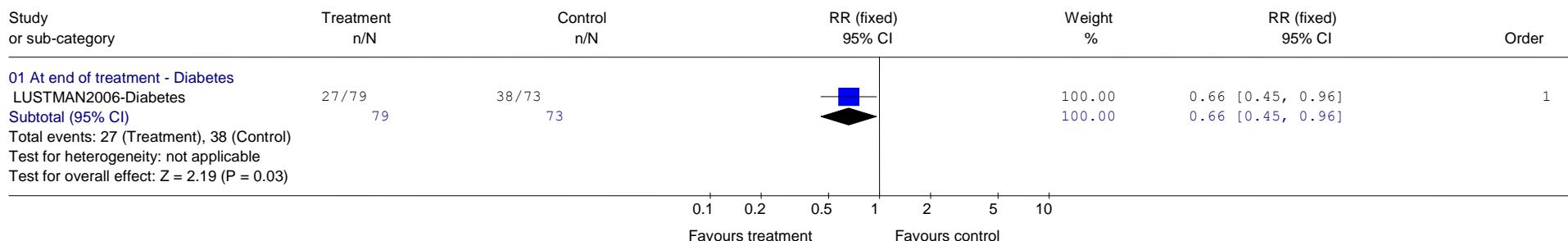
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 34 Depression: 3. Patient-rated Continuous measures



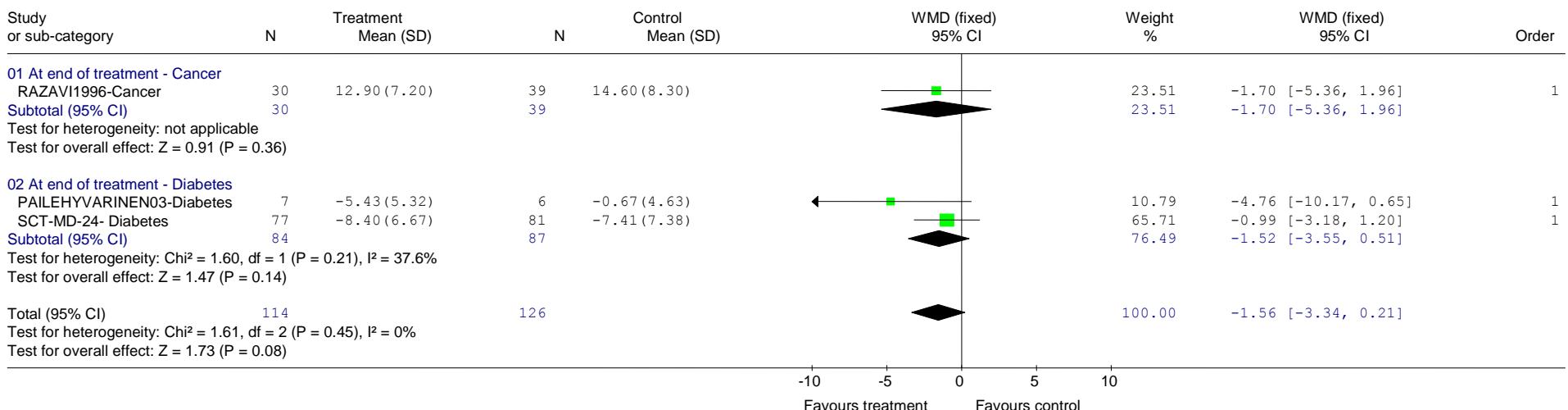
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 35 Depression: 4. Observer-rated Continuous measures



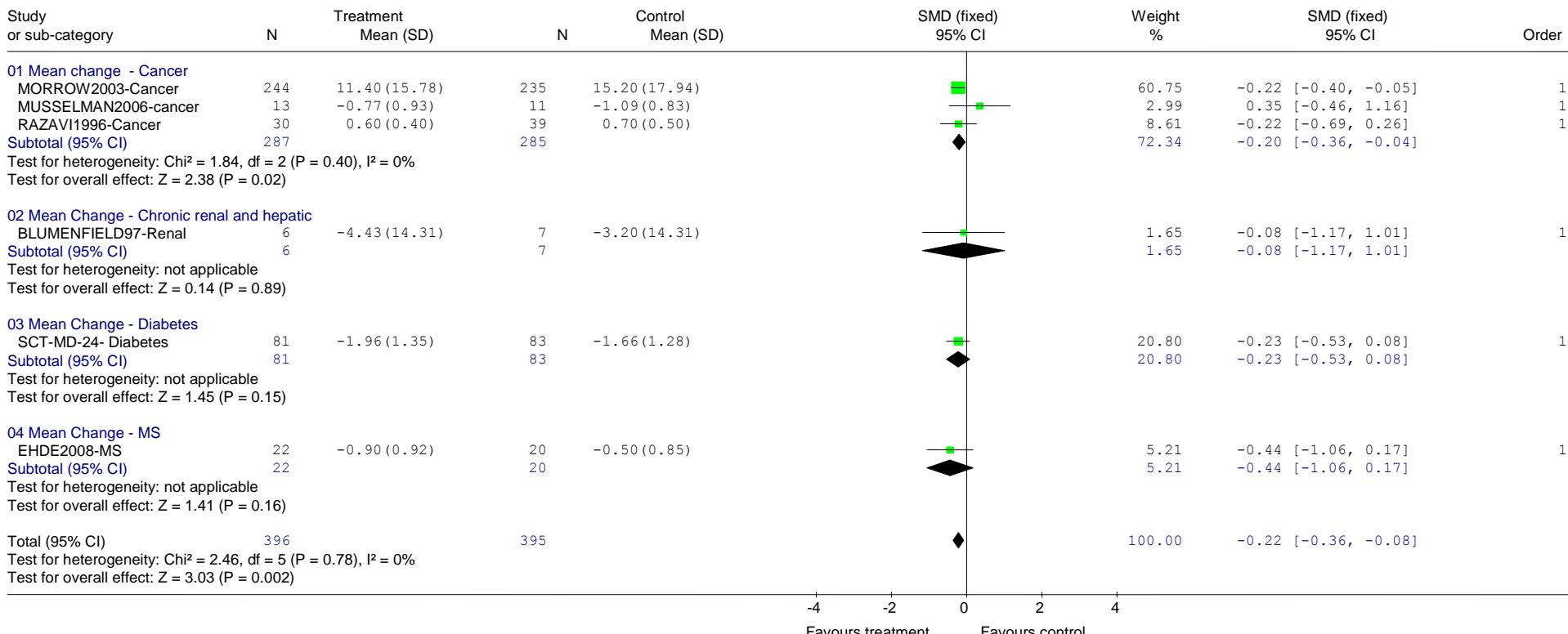
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 36 Depression: 5. Recurrence/ Relapse of depression



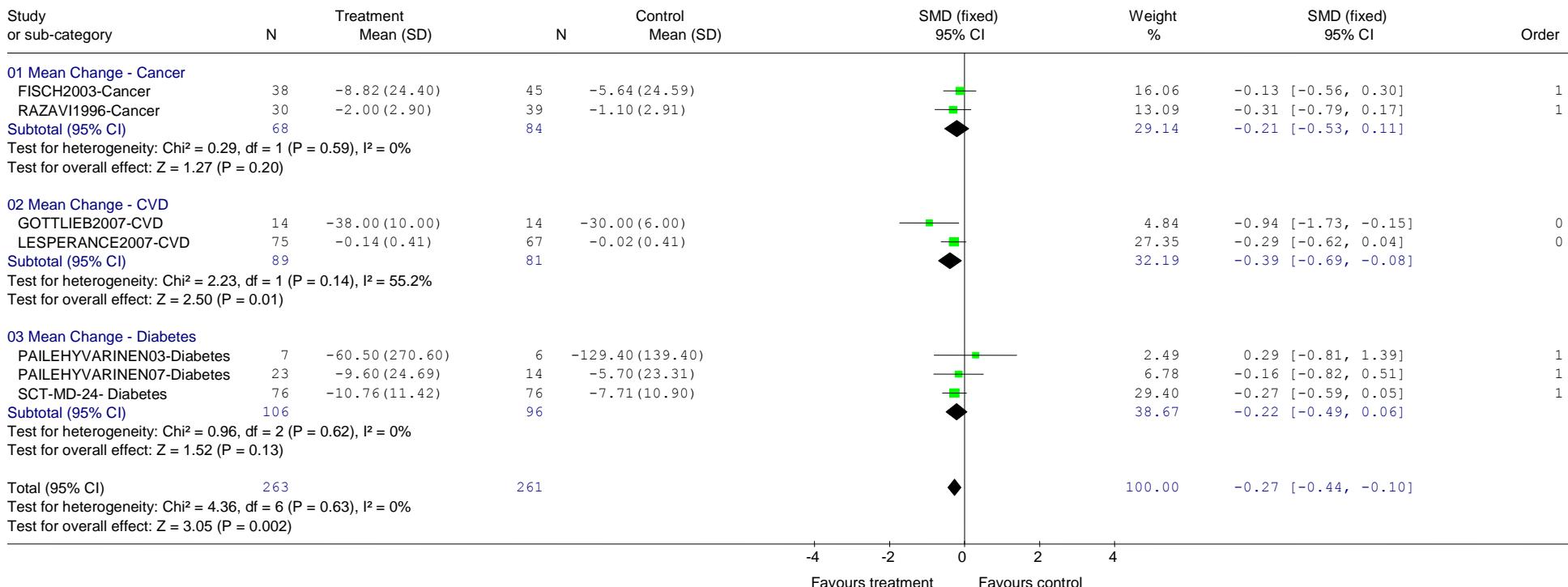
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 40 Mental State: Continuous measures 1. Hamilton Anxiety Scale



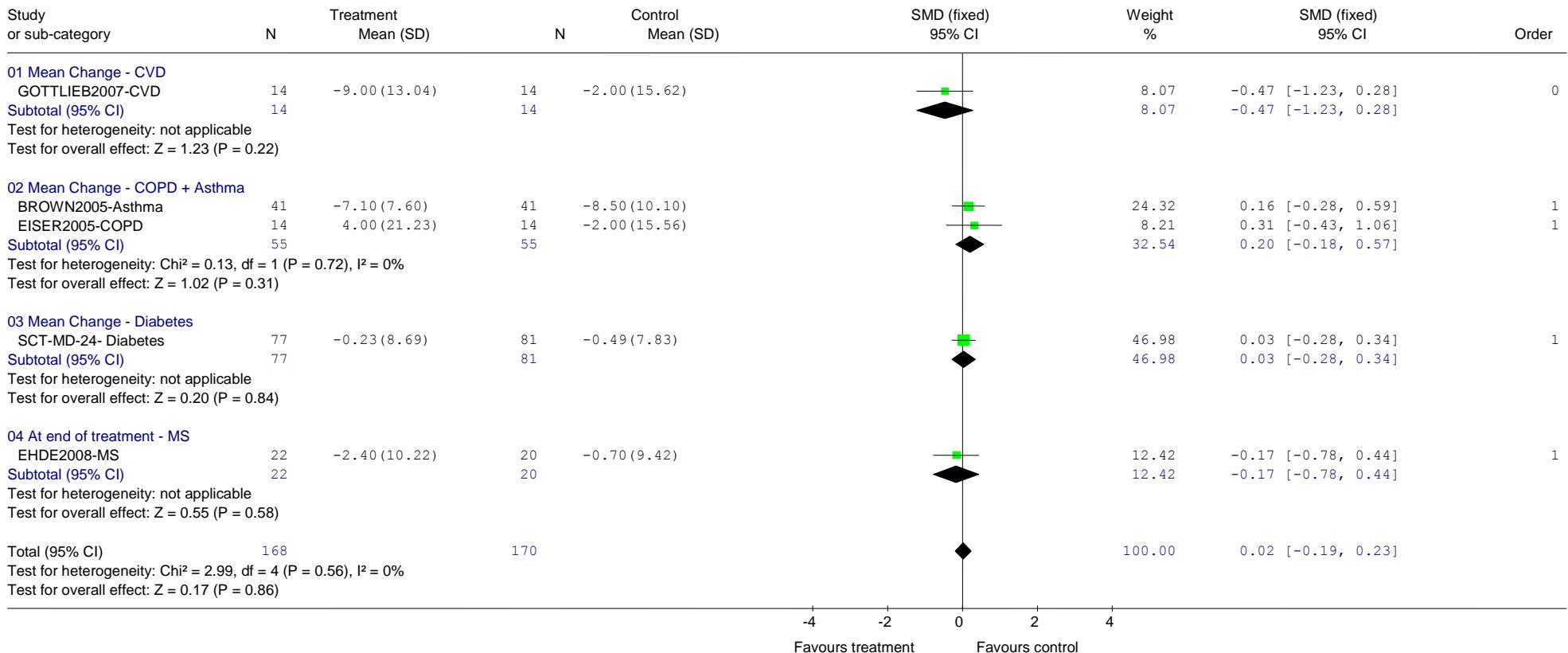
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 41 Mental State: Continuous measures 2. Global symptom measures e.g CGI, SCL-90, POMS



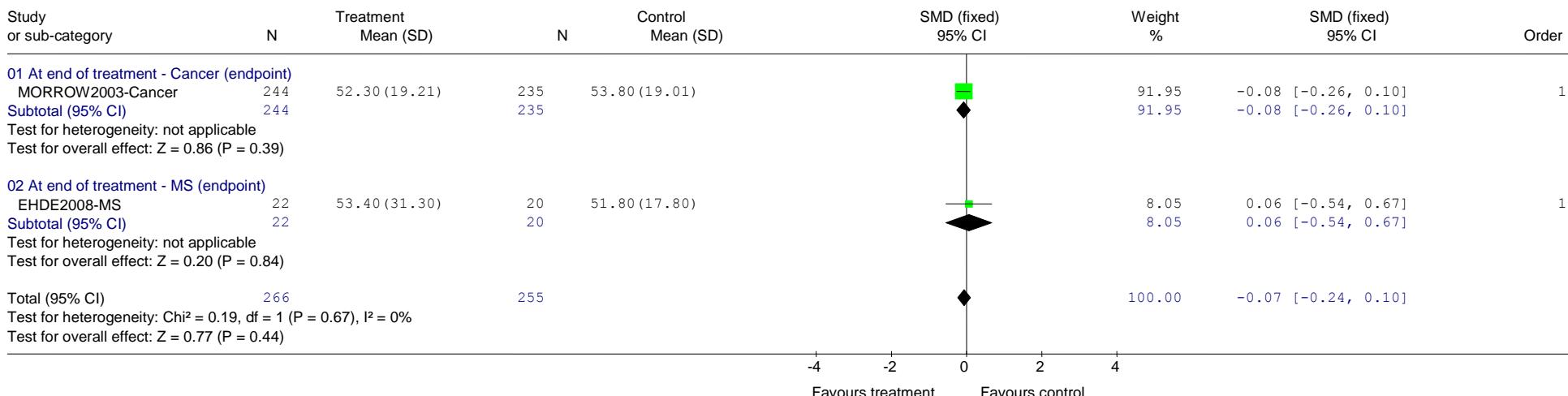
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 50 QoL: 1. Continuous measures e.g. SQOLI, FACT-G



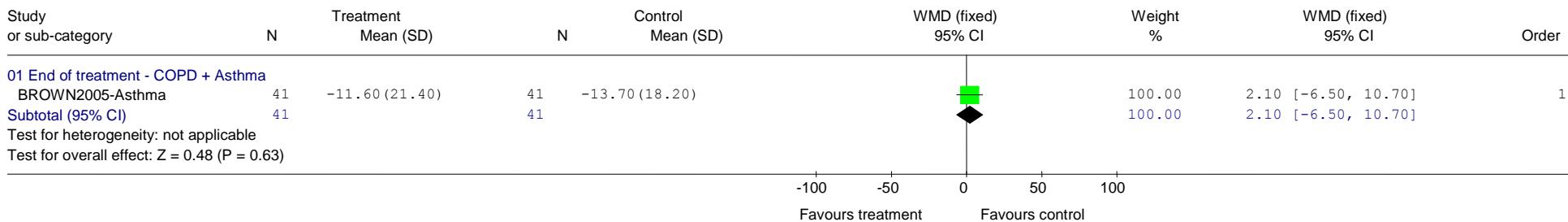
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 51 Physical outcome / QoL - General physical functioning/ well-being (SF-36 physical component)



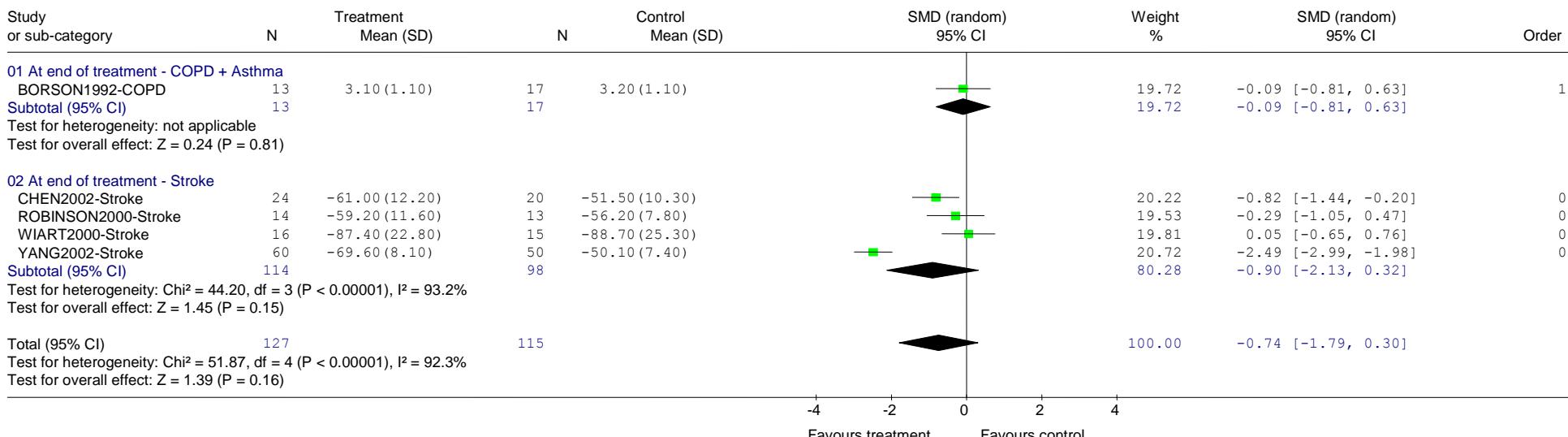
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 52 Physical health outcome. 1. Fatigue



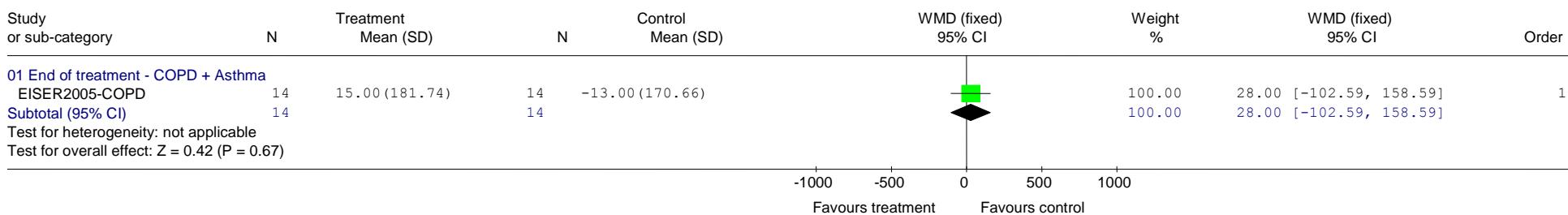
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 53 Physical health outcome: 2. Asthma Control Questionnaire



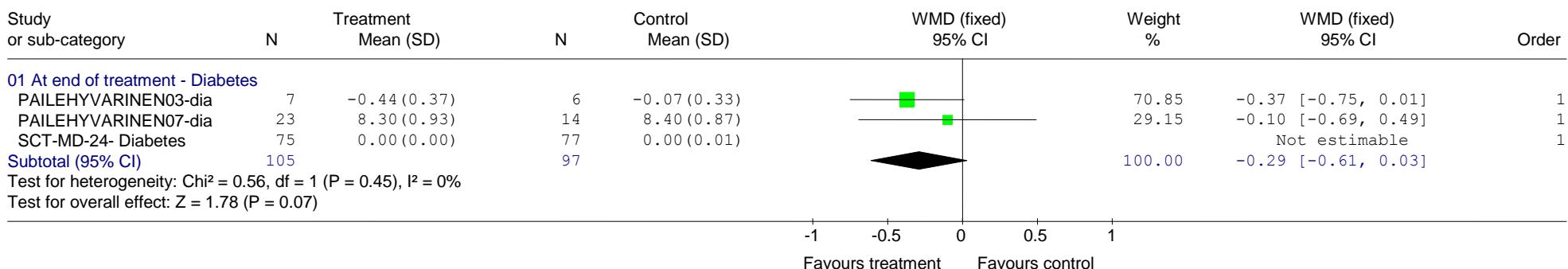
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 54 Physical health outcome: 3. Physical activity - PFSI, ADLI, functional independence



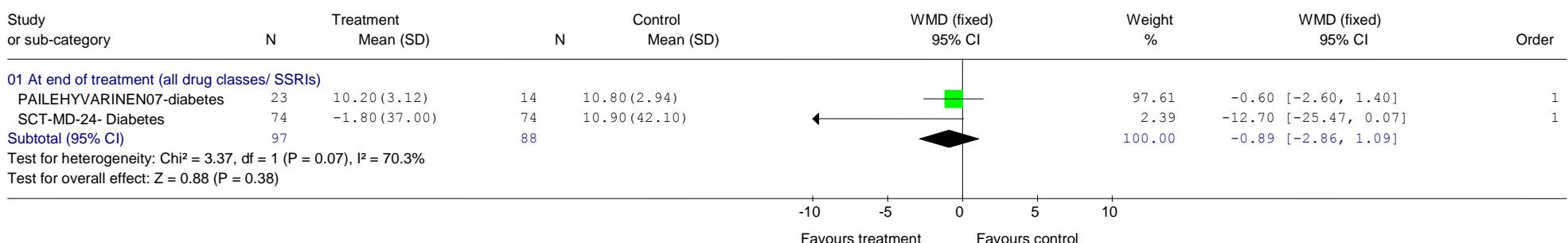
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 55 Physical health outcome: 4. Exercise tolerance test 6MWD (signs reversed)



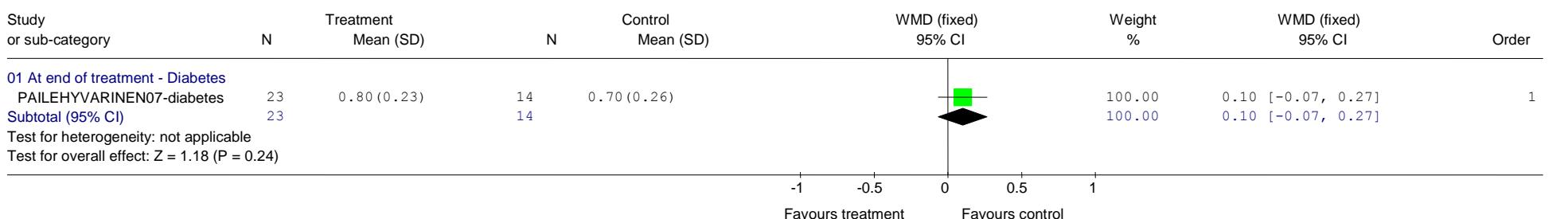
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 56 Physical health outcome: 5. GHbA1c (%)



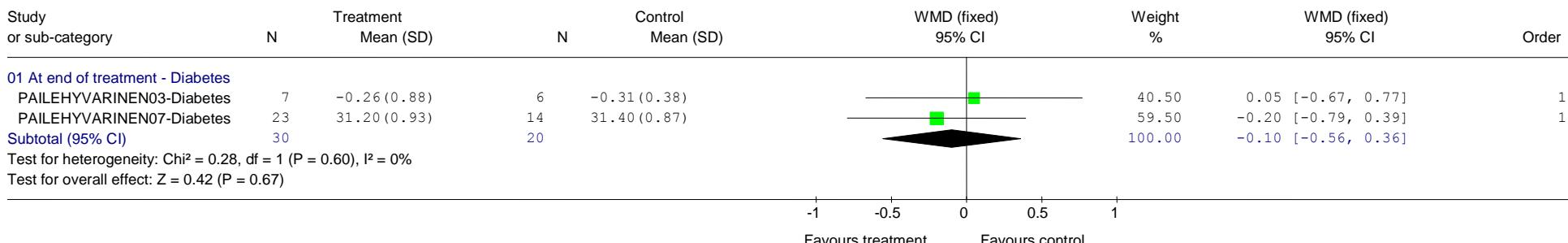
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 57 Physical health outcome: 6. Serum Glucose (mmol/l)



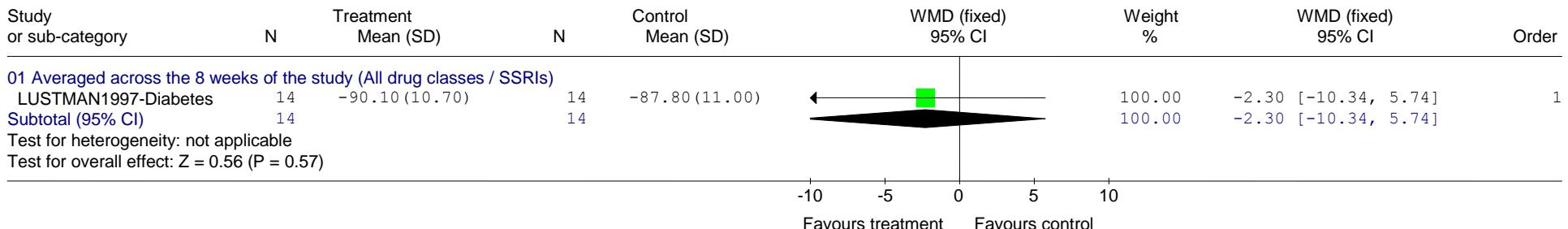
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 58 Physical health outcome. 7. Serum C-peptide (nmol/l)



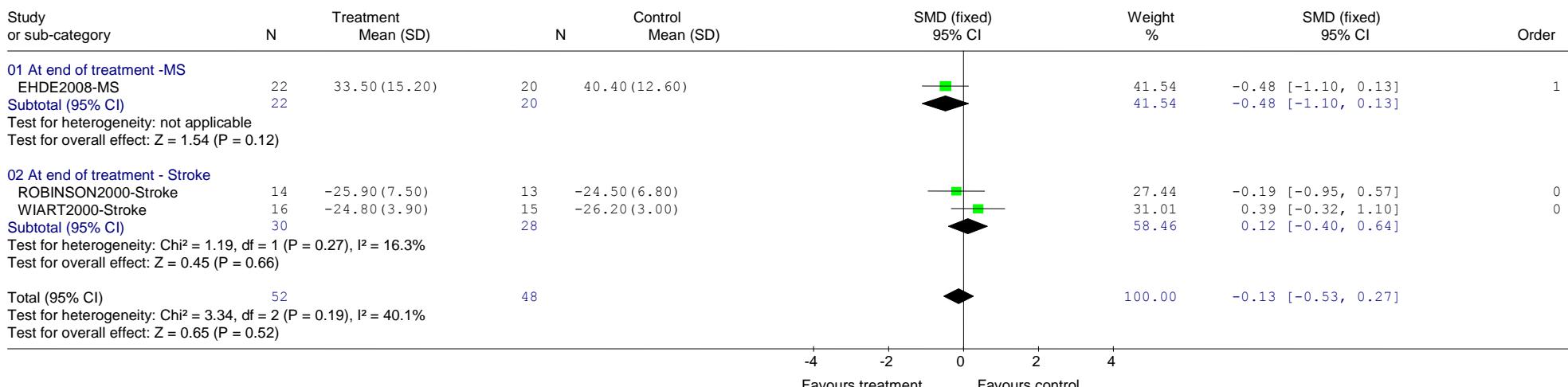
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 59 Physical health outcome: 8. BMI



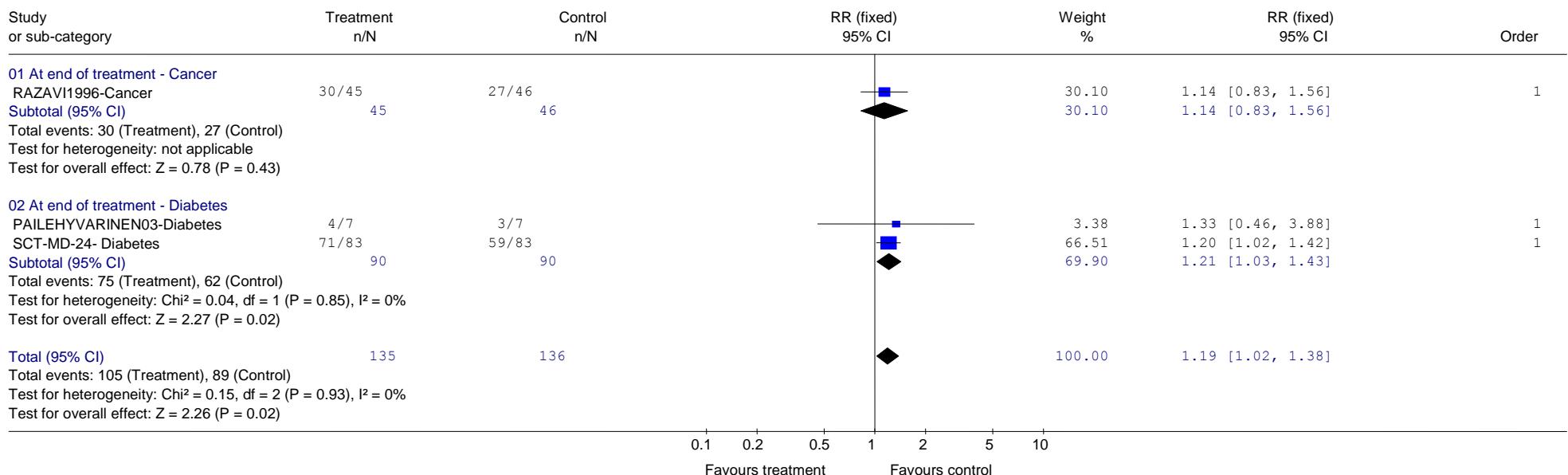
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 60 Physical health outcome: 9. Medication compliance



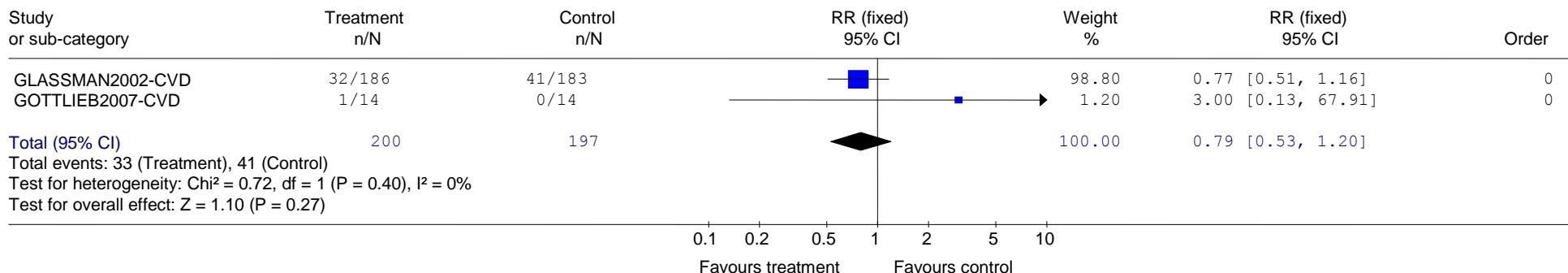
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 61 Physical health outcome: 10. Cognitive deficits (PDQ, MMSE)



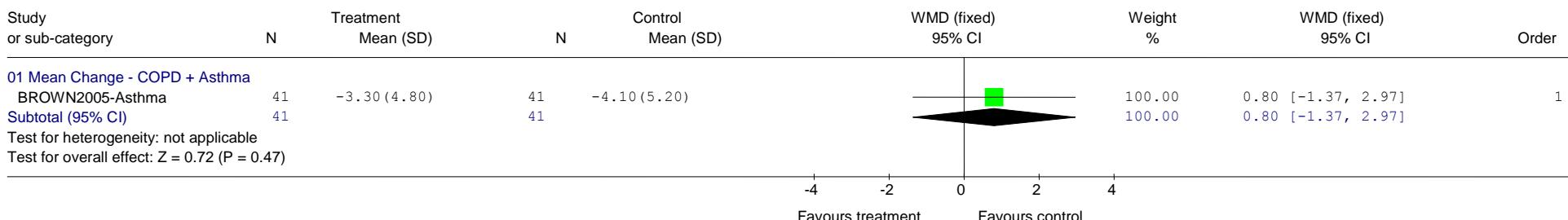
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 70 AE: 1. Reported at least 1 adverse event



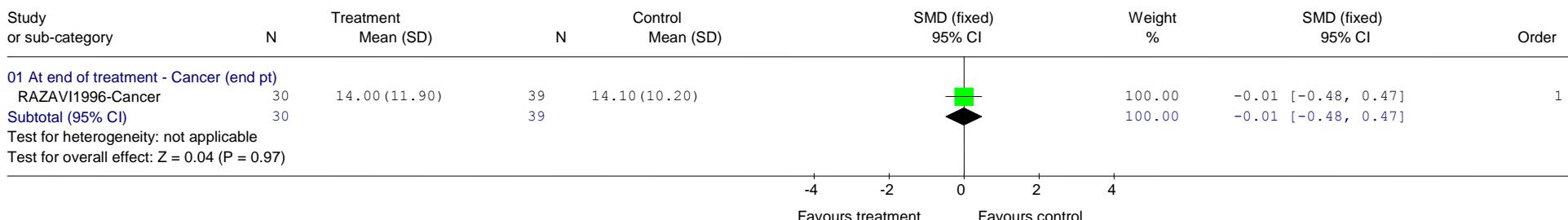
Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 71 AE: 2. Death or cardiac rehospitalisation



Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 72 AE: 3. Continuous measures PRD-III

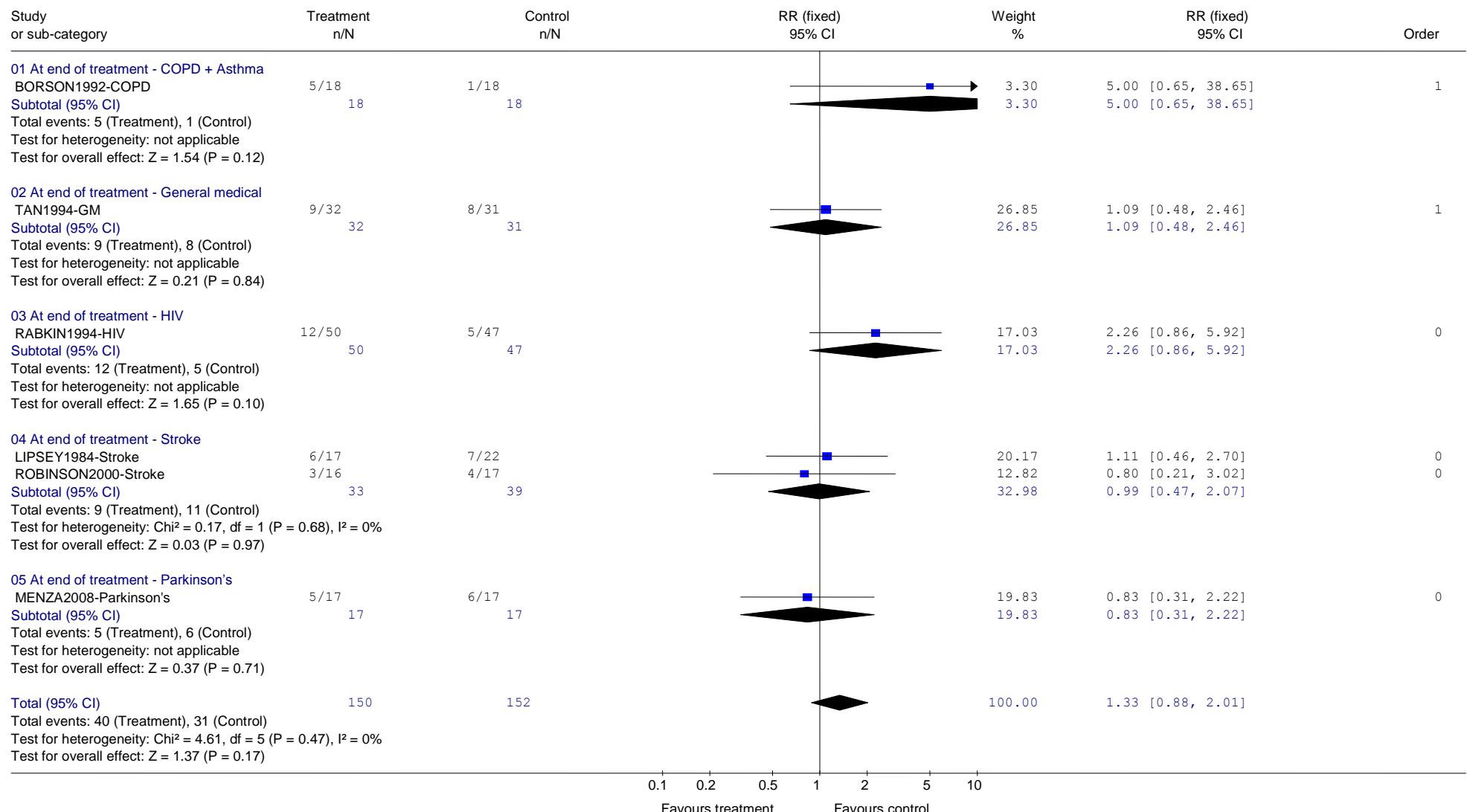


Review: DCHP - Pharm - by drug class  
 Comparison: 02 SSRIs versus placebo  
 Outcome: 73 Treatment tolerance: 1. Observer-rated continuous measures

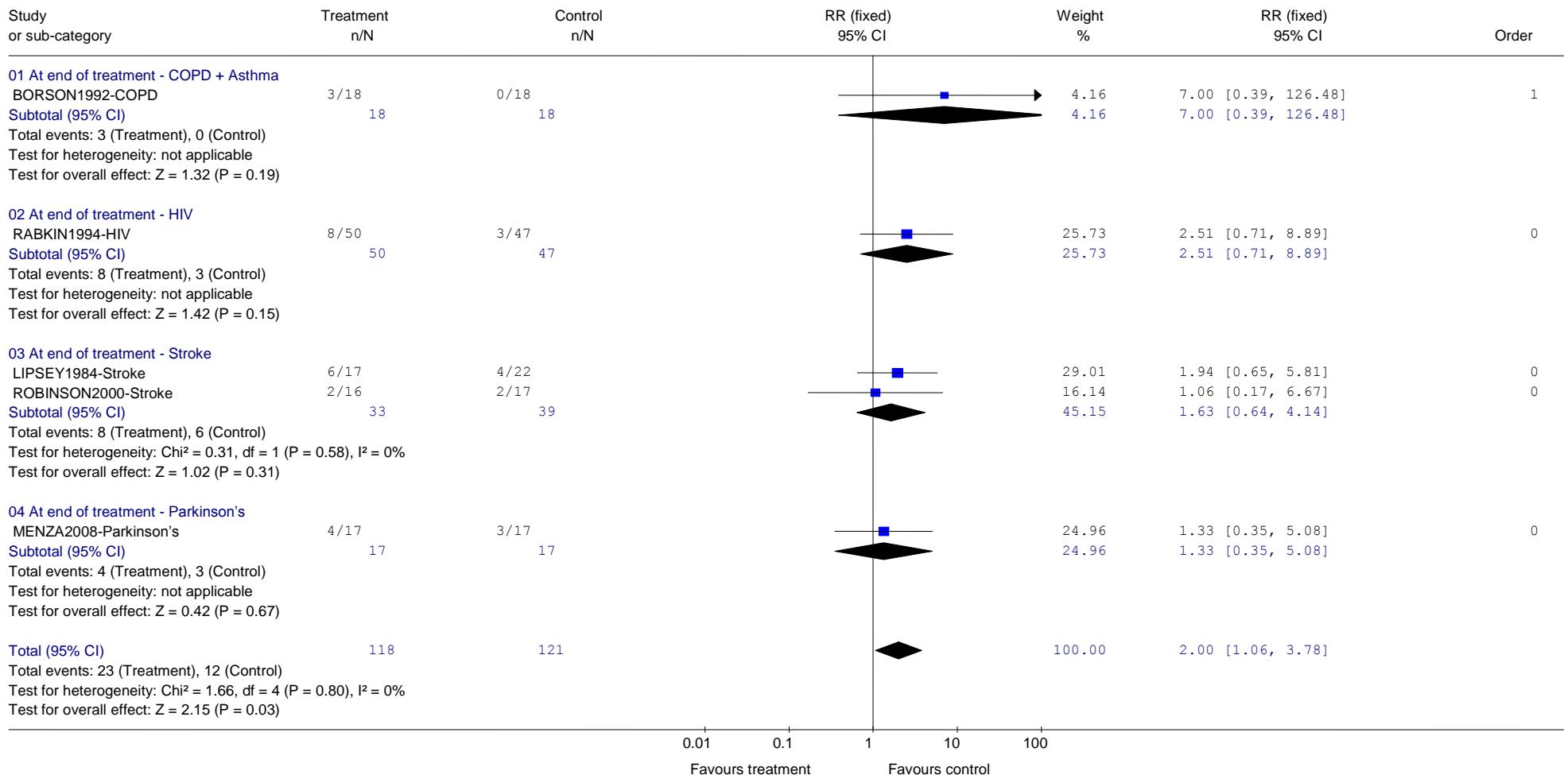


## TCAs versus placebo

Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 01 Leaving the study early: Any reason



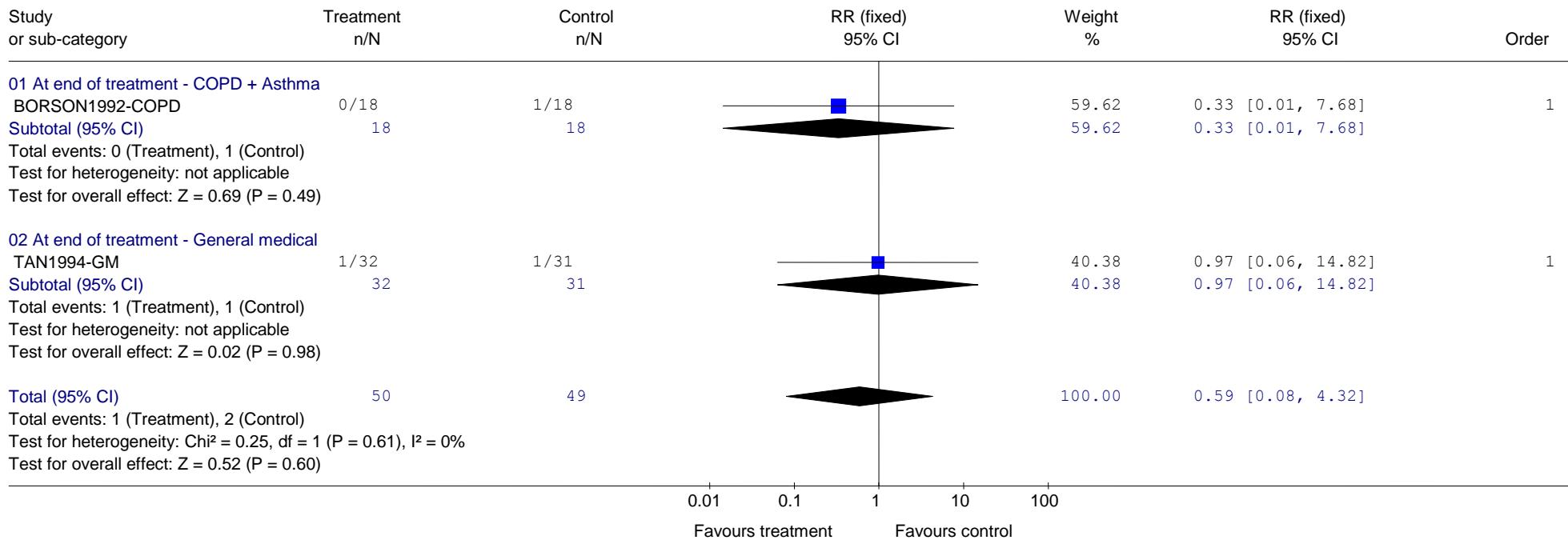
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 02 Leaving due to adverse events



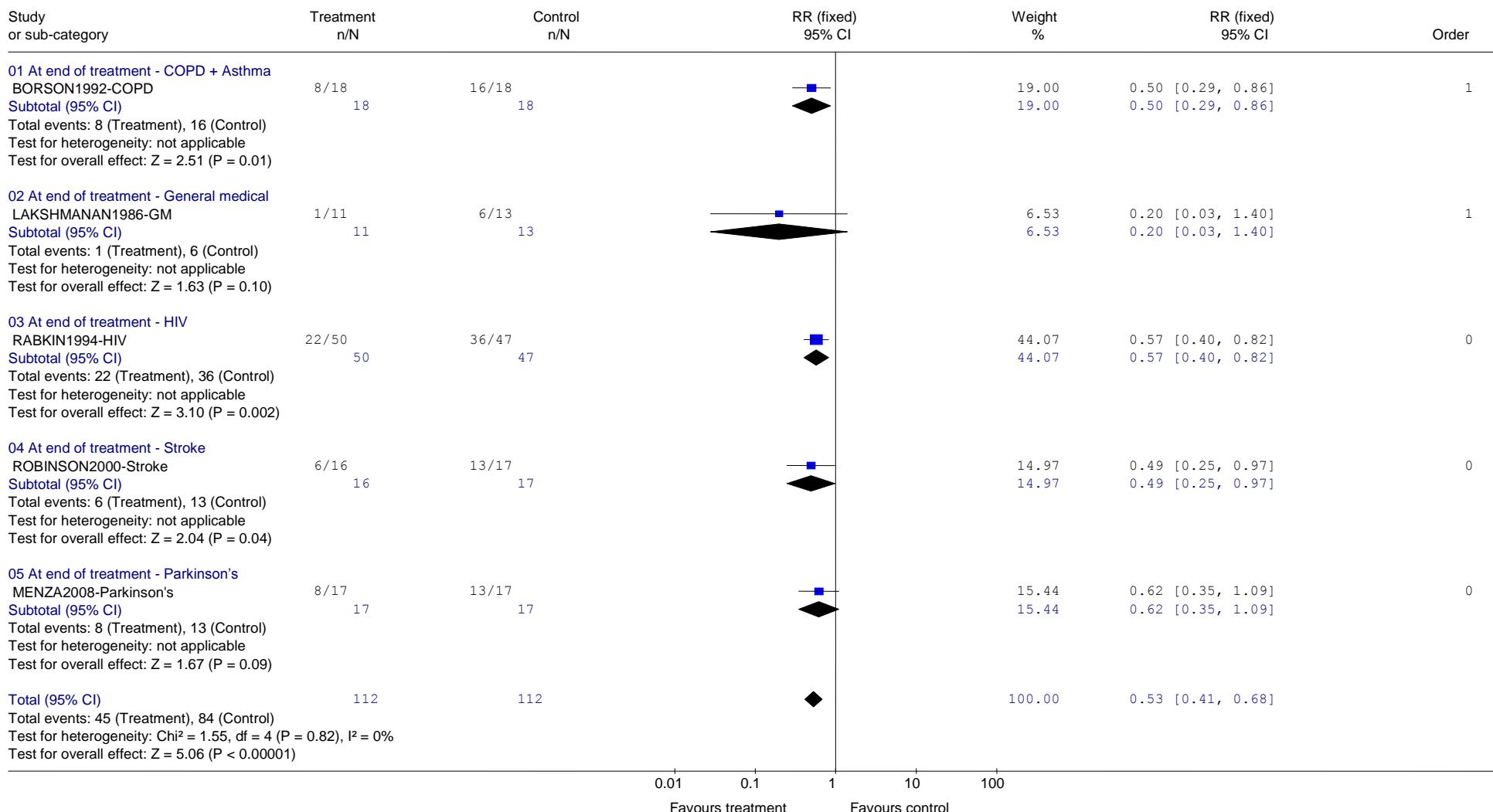
Review: DCHP - Pharm - by drug class

Comparison: 03 TCAs versus placebo

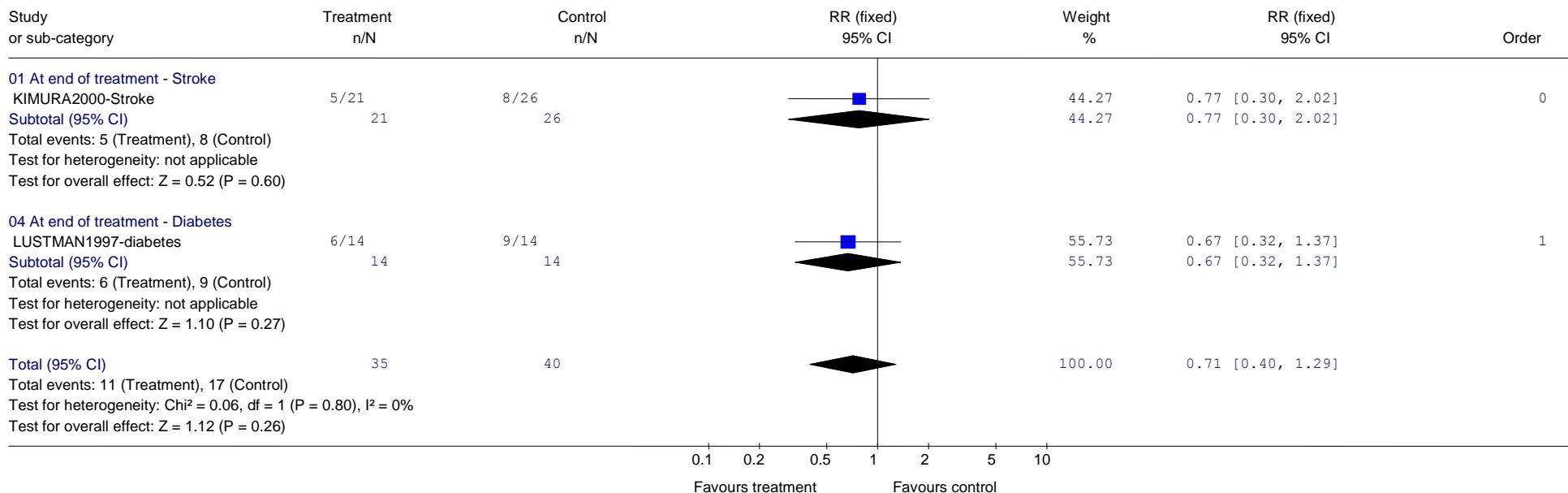
Outcome: 05 Mortality



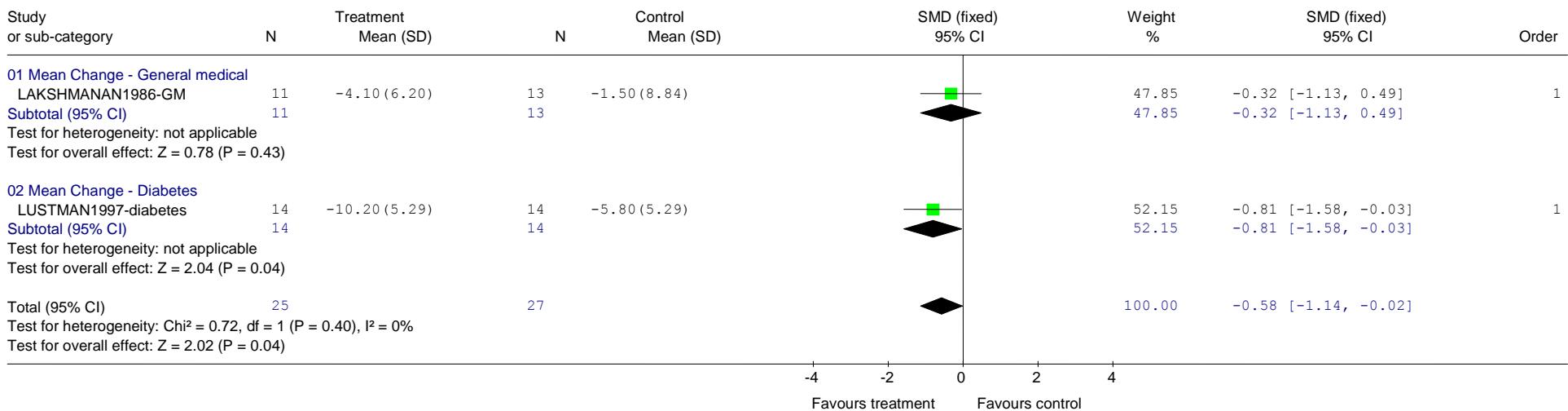
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 10 Depression: 1. Non-response (<50% improvement) - observer rated



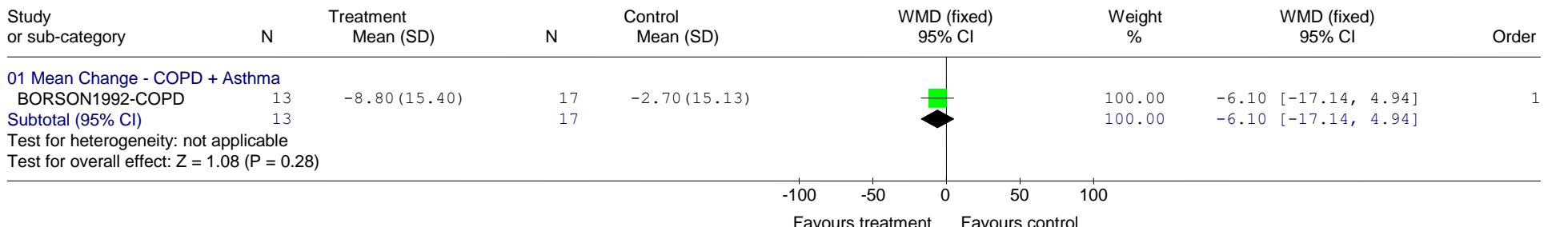
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 11 Depression: 2. Not achieving success/ remission (reaching a specified cut-off) Patient-rated



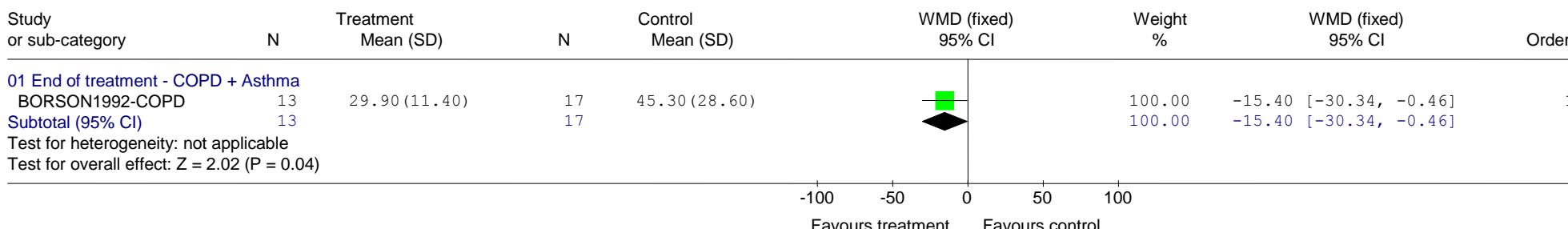
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 12 Depression: 3. Patient-rated continuous measures (End of treatment)



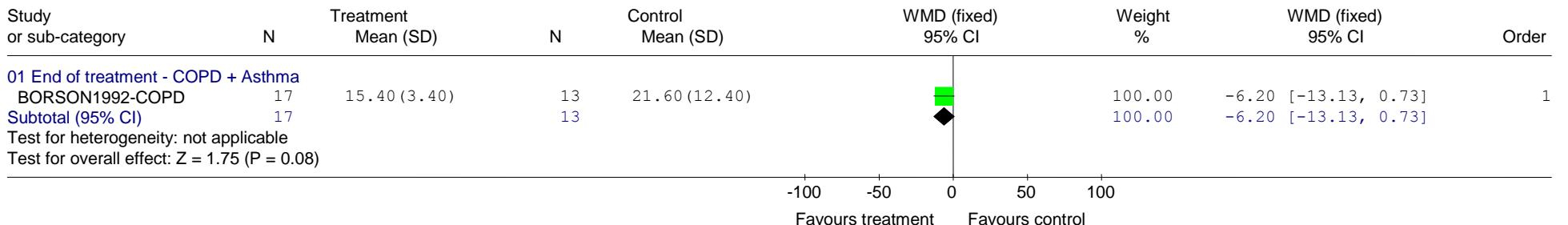
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 20 Physical outcome / QoL - General physical functioning/ well-being (SF-36 etc)



Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 25 Physical health outcome: 1. Symptom anxiety PRAS



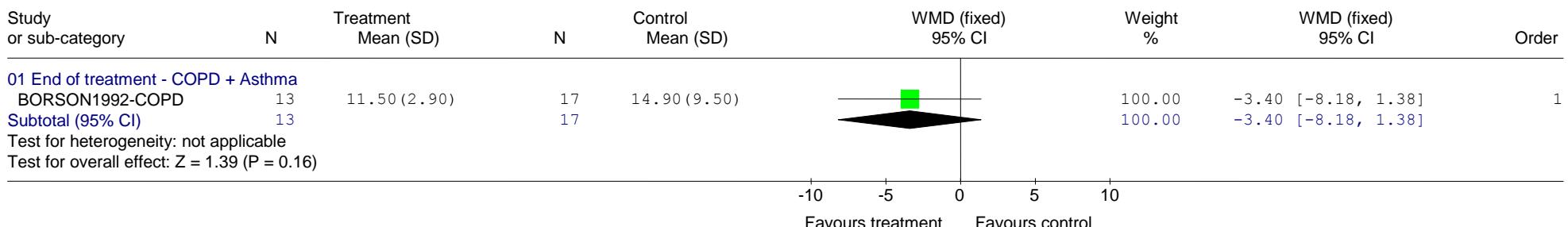
Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 26 Physical health outcome: 2. Somatic symptoms



Review: DCHP - Pharm - by drug class

Comparison: 03 TCAs versus placebo

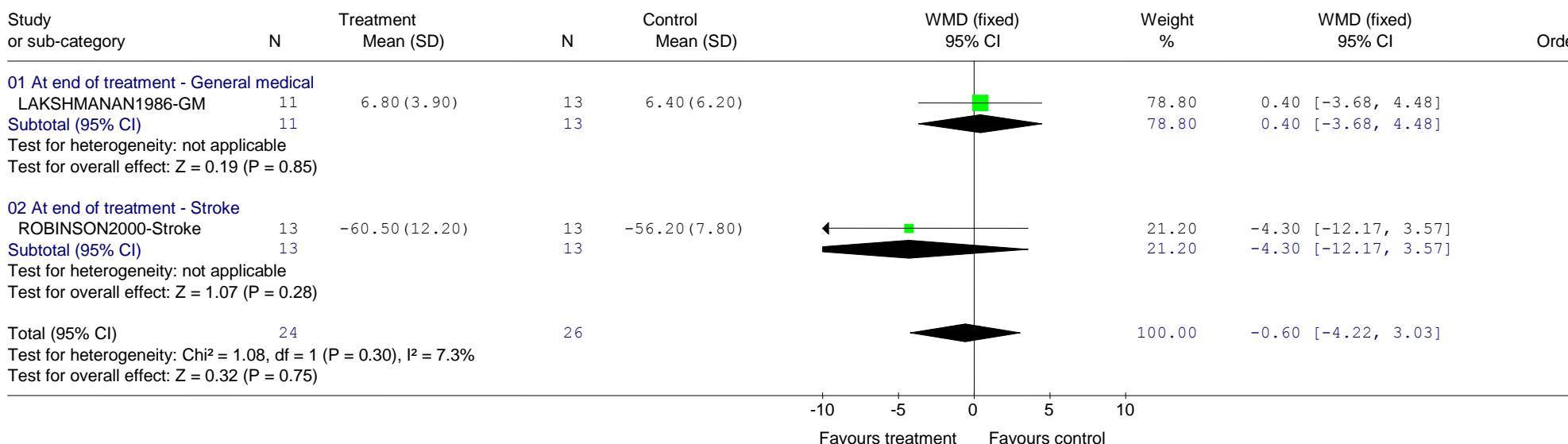
Outcome: 27 Physical health outcome: 3. Symptoms associated with breathing



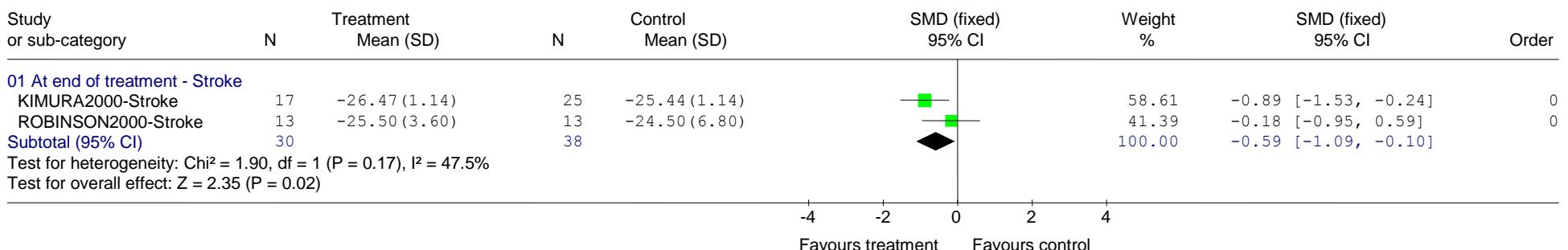
Review: DCHP - Pharm - by drug class

Comparison: 03 TCAs versus placebo

Outcome: 28 Physical health outcome: 4. Physical activity (HVID, ADL)

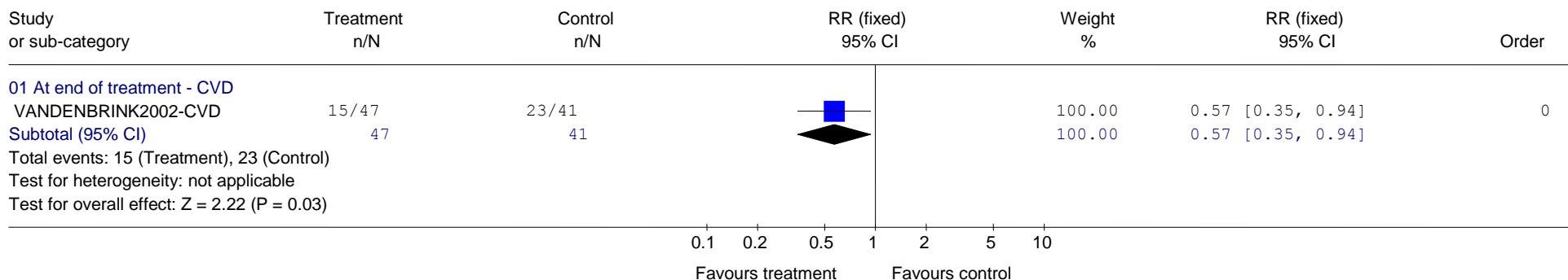


Review: DCHP - Pharm - by drug class  
 Comparison: 03 TCAs versus placebo  
 Outcome: 29 Physical health outcome: 5. Cognitive function (MMSE)

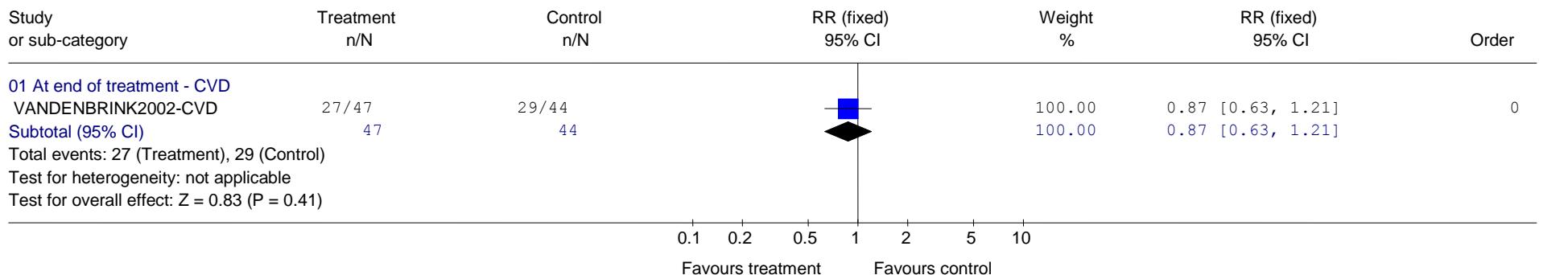


## Third-generation drugs versus placebo

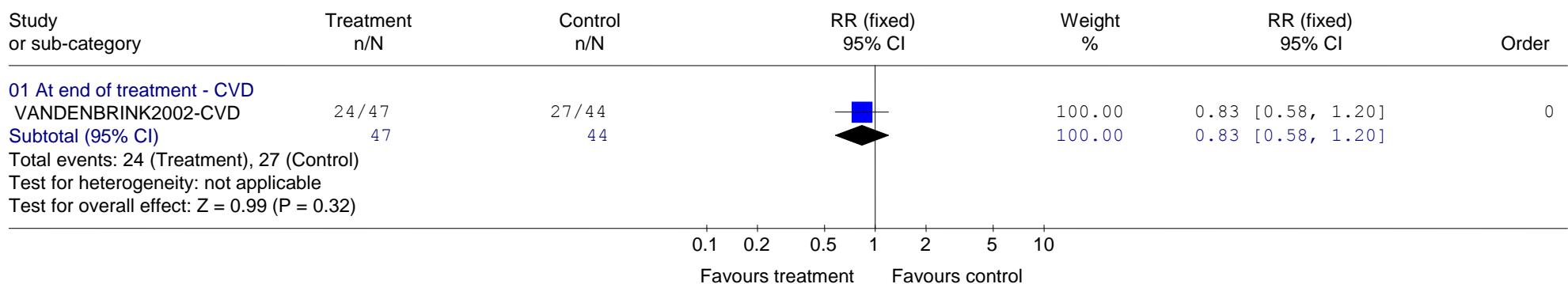
Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 01 Leaving the study early: any reason



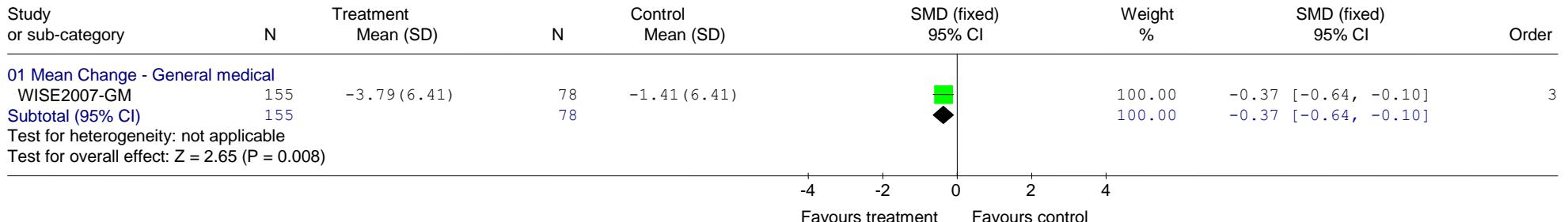
Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 21 Depression: 1. Not achieving success/ remission (reaching a specified cut-off) - observer rated



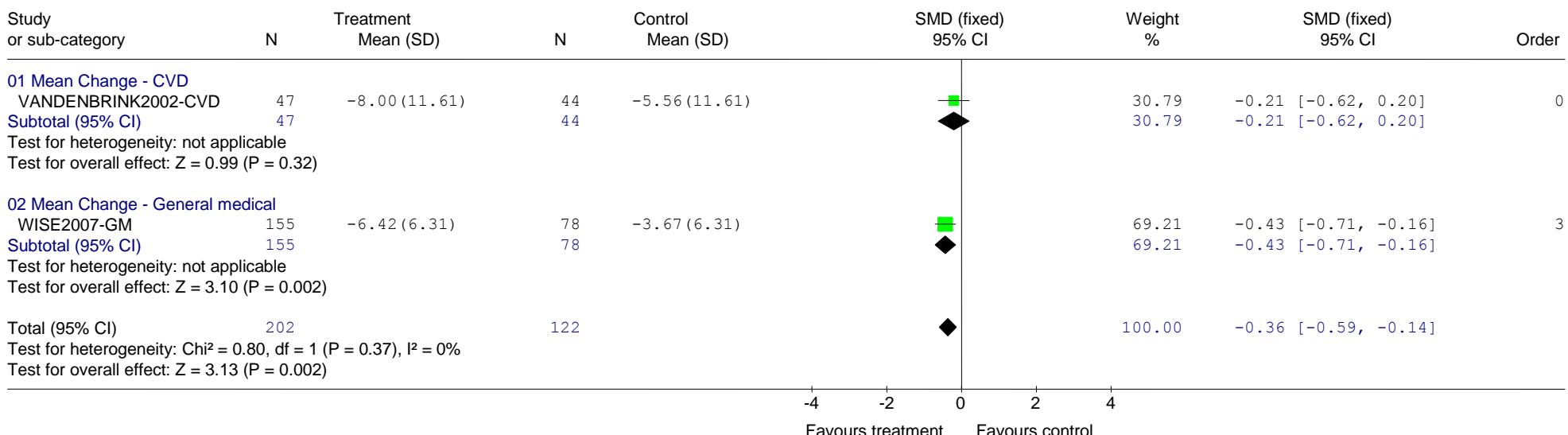
Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 22 Depression: 2. Non-response (<50% reduction) - observer rated



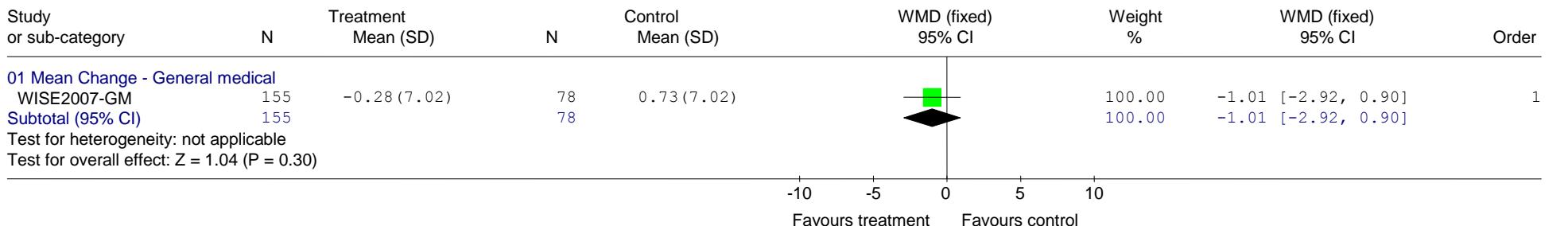
Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 23 Depression 3: Continuous measures patient rated, e.g, BDI, HADS GDS



Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 24 Depression: 4. Observer-rated continuous measures



Review: DCHP - Pharm - by drug class  
 Comparison: 04 3rd generation ADs versus placebo  
 Outcome: 30 Physical outcome / QoL - General physical functioning/ well-being (SF-36 etc)

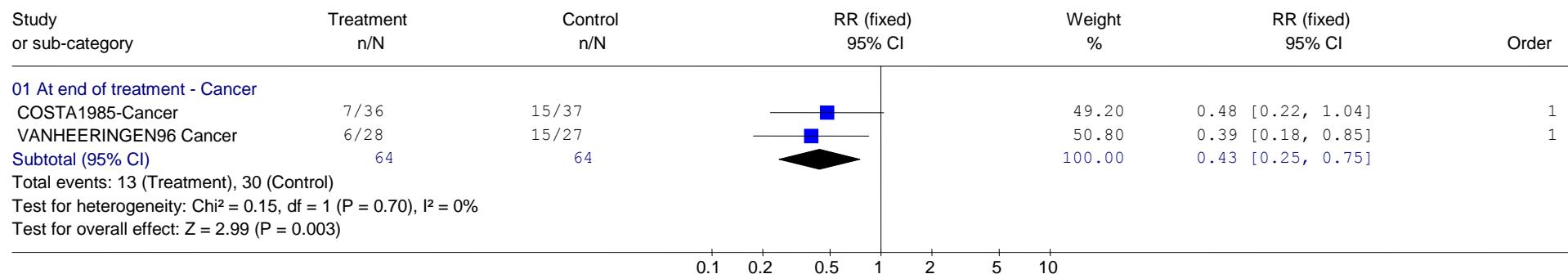


## Mianserin versus placebo

Review: DCHP - Pharm - by drug class

Comparison: 05 Mianserin versus placebo

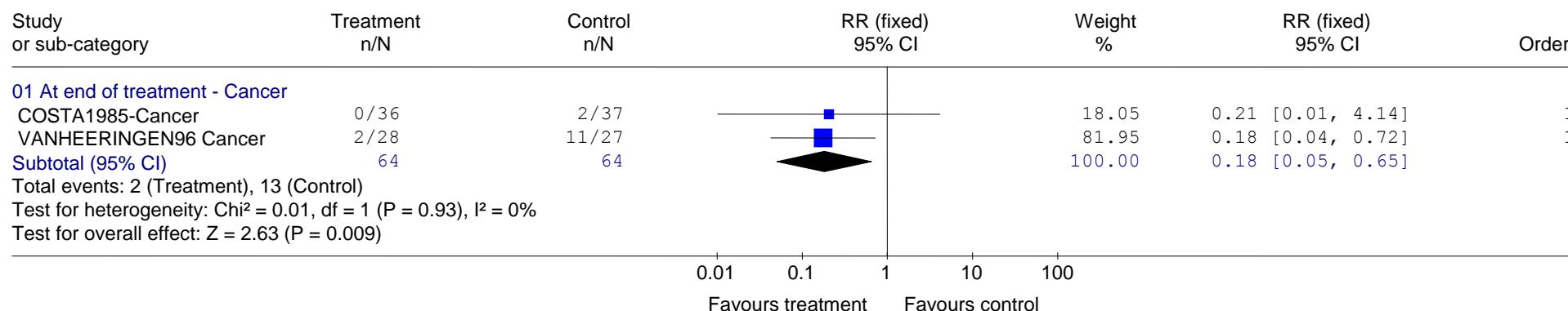
Outcome: 01 Leaving the study early: Any reason



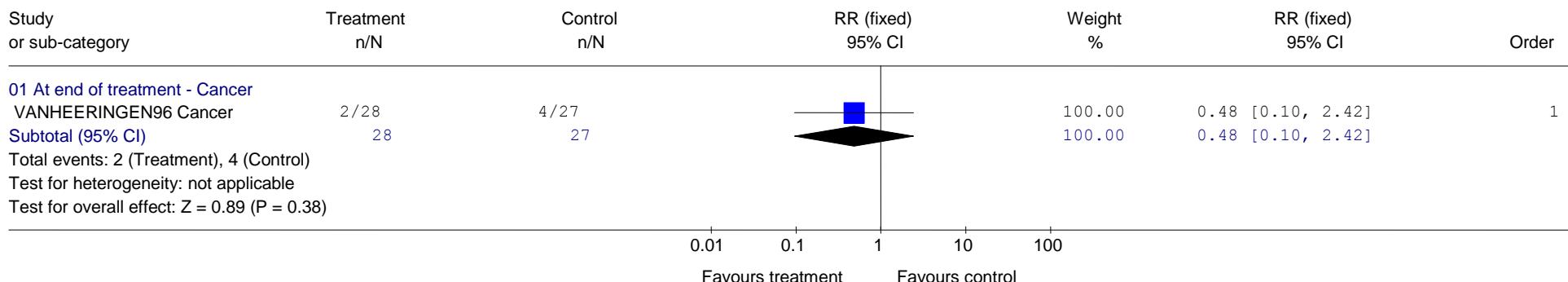
Review: DCHP - Pharm - by drug class

Comparison: 05 Mianserin versus placebo

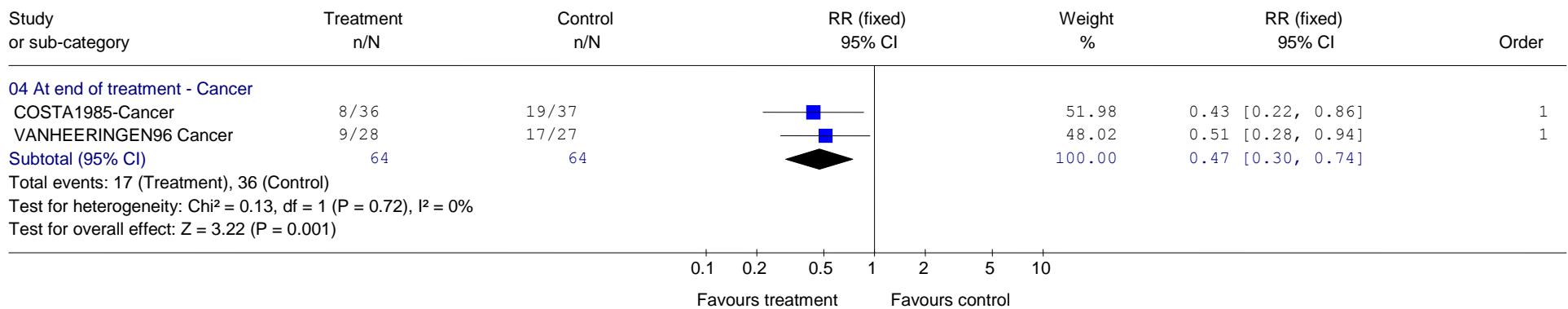
Outcome: 02 Leaving the study early due to lack of efficacy



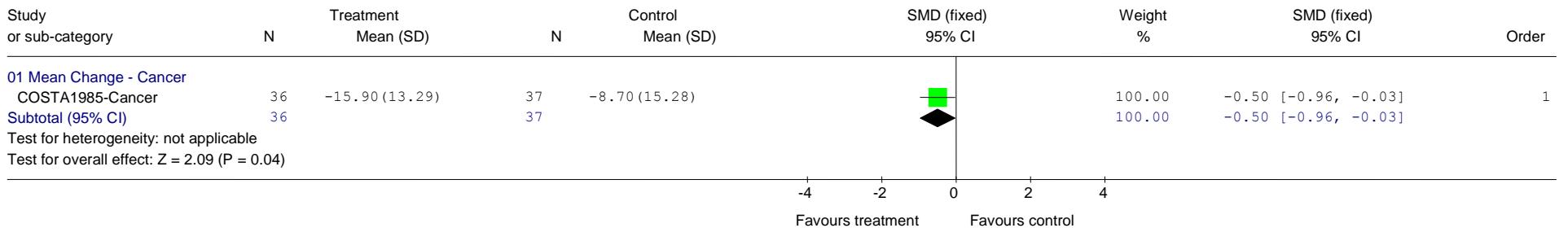
Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin versus placebo  
 Outcome: 03 Leaving the Study early: Due to adverse events



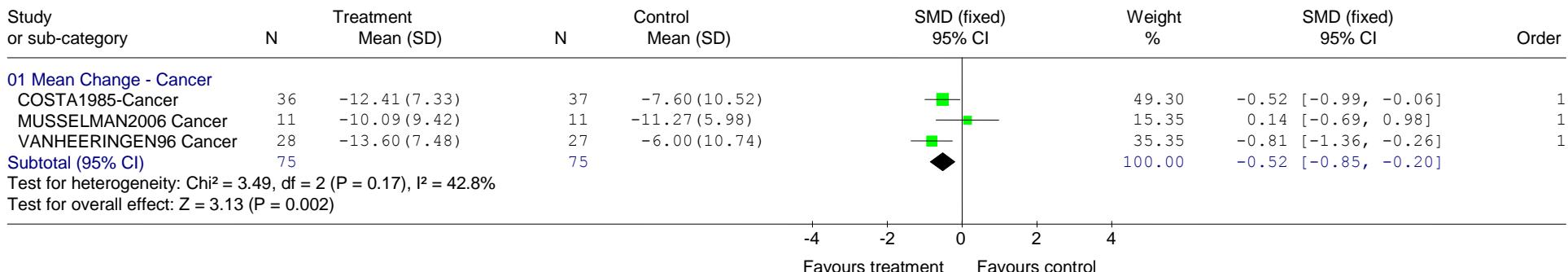
Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin versus placebo  
 Outcome: 04 Depression: 2. Non-response (not achieving 50% reduction from baseline) - observer rated e.g. HAMD, MADRS



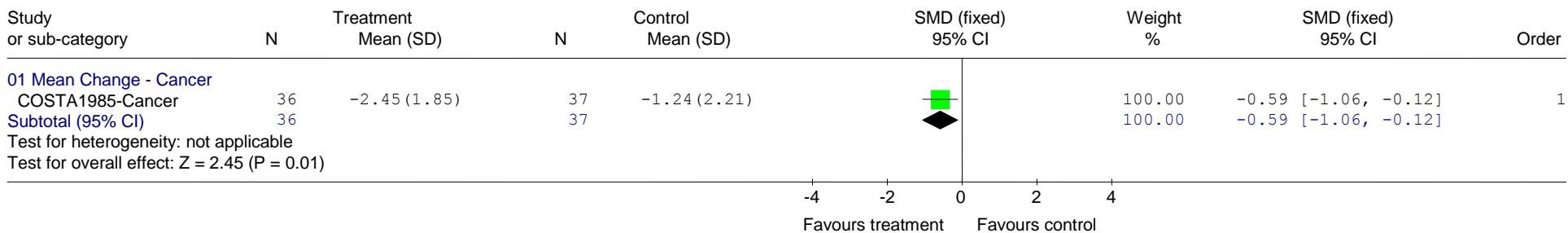
Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin versus placebo  
 Outcome: 05 Depression: 3. Patient-rated continuous measures



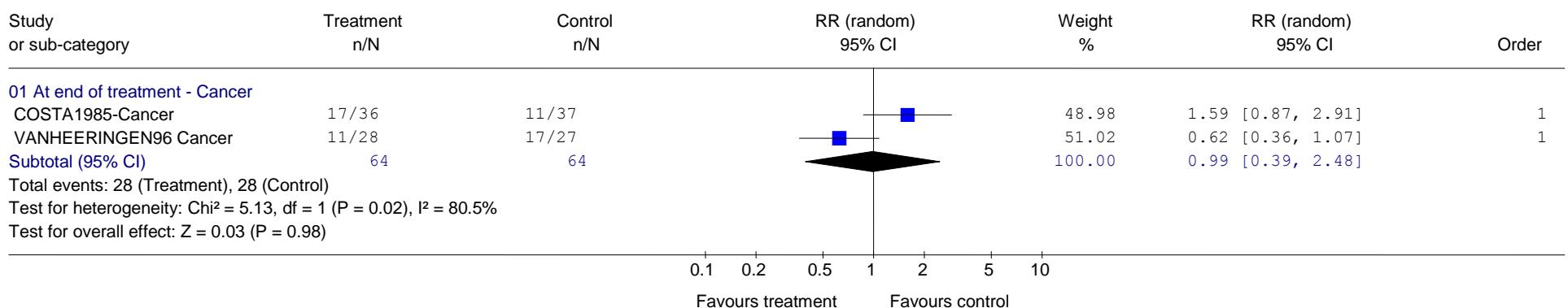
Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin vs. placebo  
 Outcome: 06 Depression: 4. Observer-rated continuous measures



Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin versus placebo  
 Outcome: 07 Mental State: Continuous measures 2. Global symptom measures e.g CGI, SCL-90, POMS



Review: DCHP - Pharm - by drug class  
 Comparison: 05 Mianserin versus placebo  
 Outcome: 08 AE: 1. Reported at least 1 adverse event

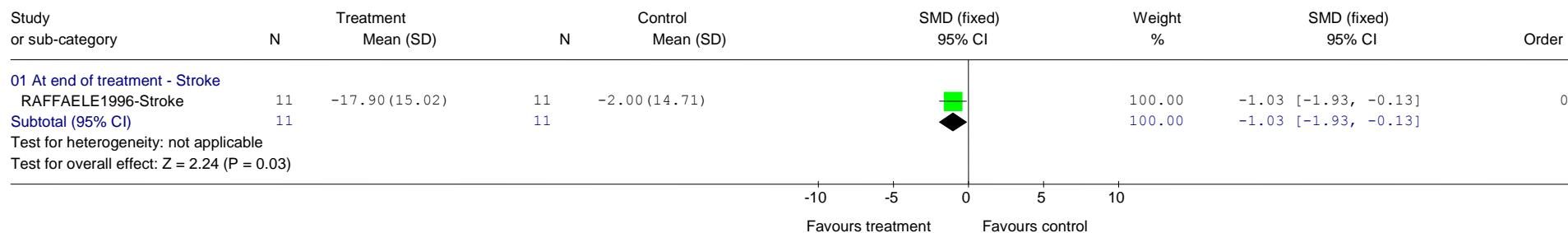


## Trazodone versus placebo

Review: DCHP - Pharm - by drug class

Comparison: 06 Trazodone versus placebo

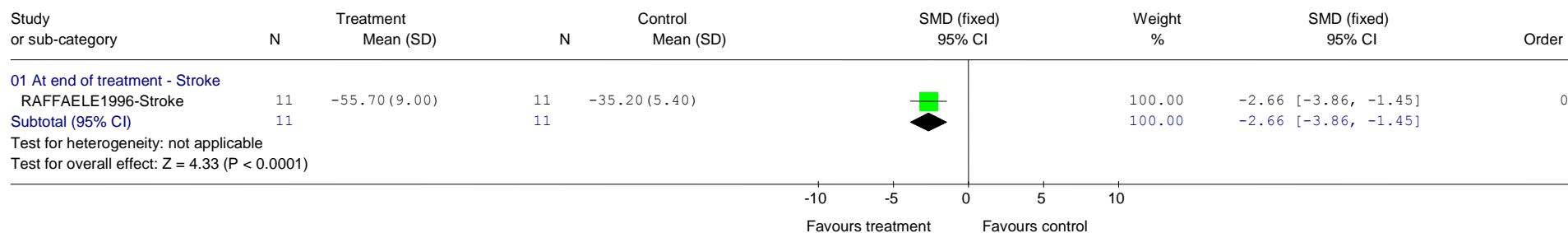
Outcome: 01 Depression: 1. Continuous measures - patient rated



Review: DCHP - Pharm - by drug class

Comparison: 06 Trazodone versus placebo

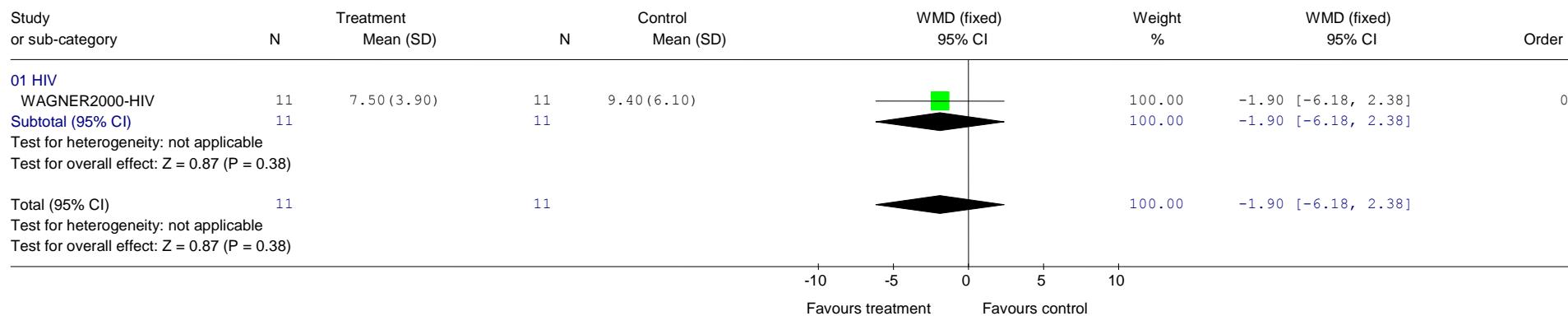
Outcome: 02 Physical health outcome: 1. Physical activities (ADL)



Review: DCHP - Pharm - by drug class

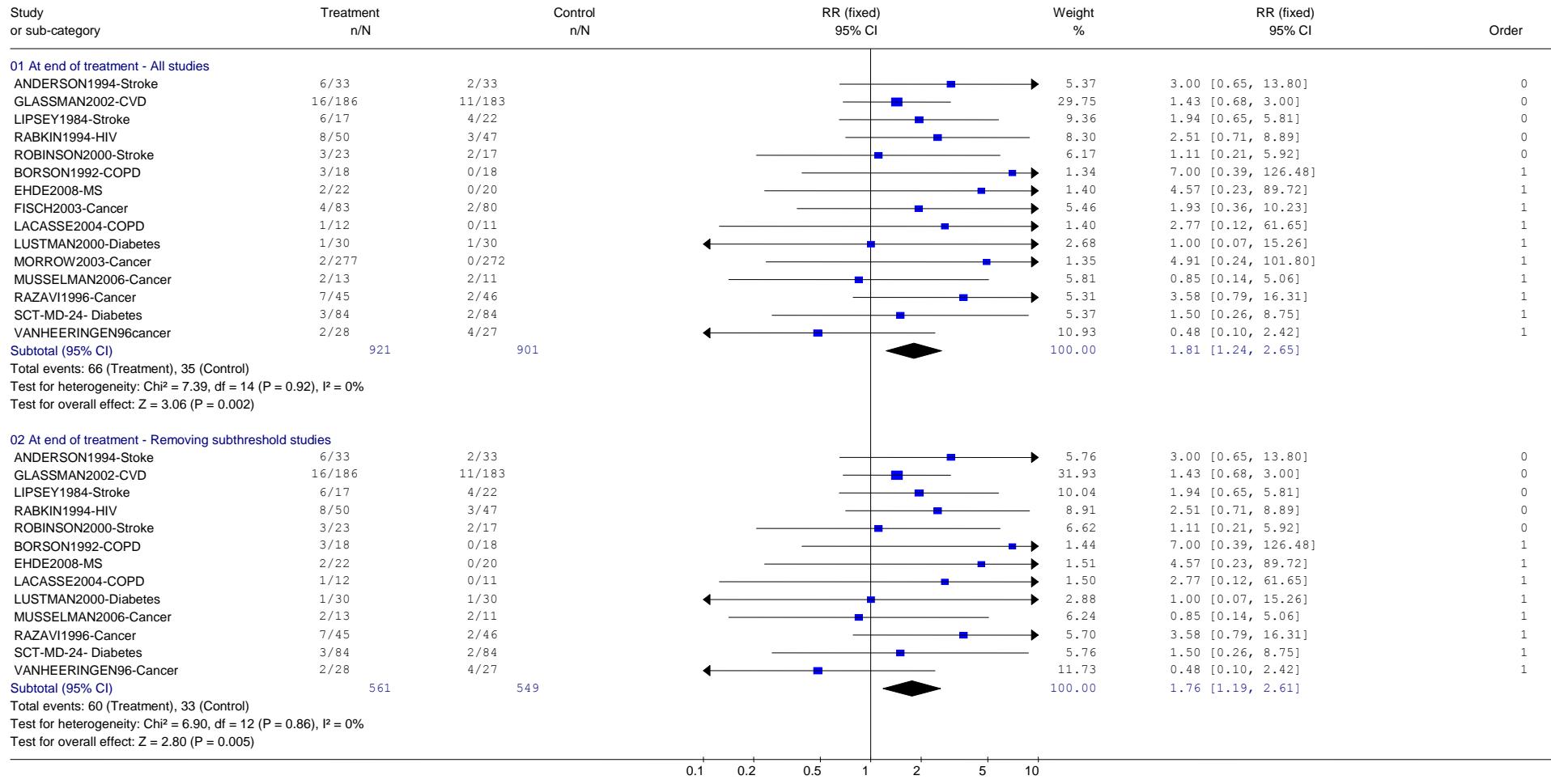
Comparison: 07 Psychostimulants versus placebo

Outcome: 01 Mean depression

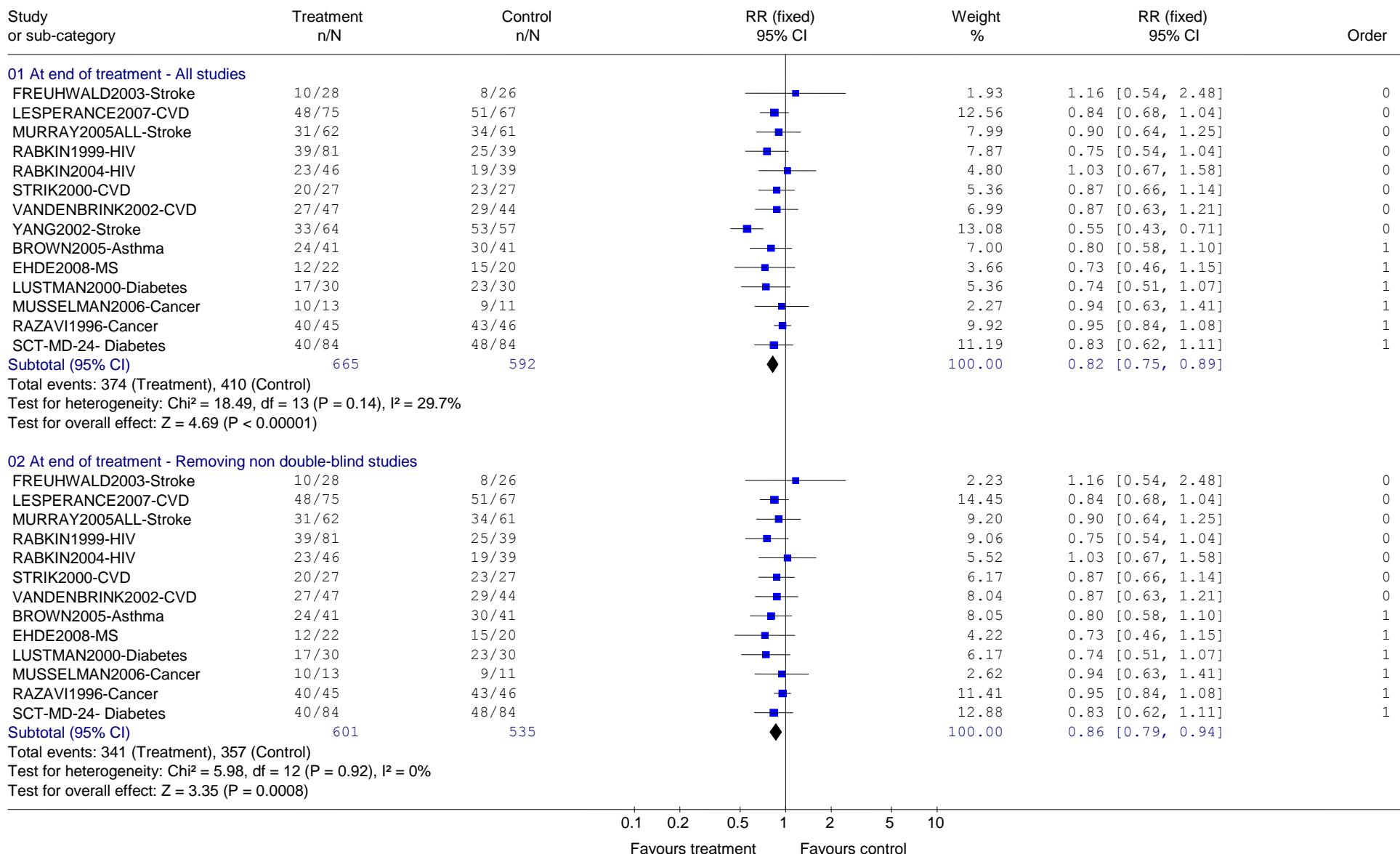


## Any antidepressant versus placebo – sensitivity analysis removing non-blinded studies

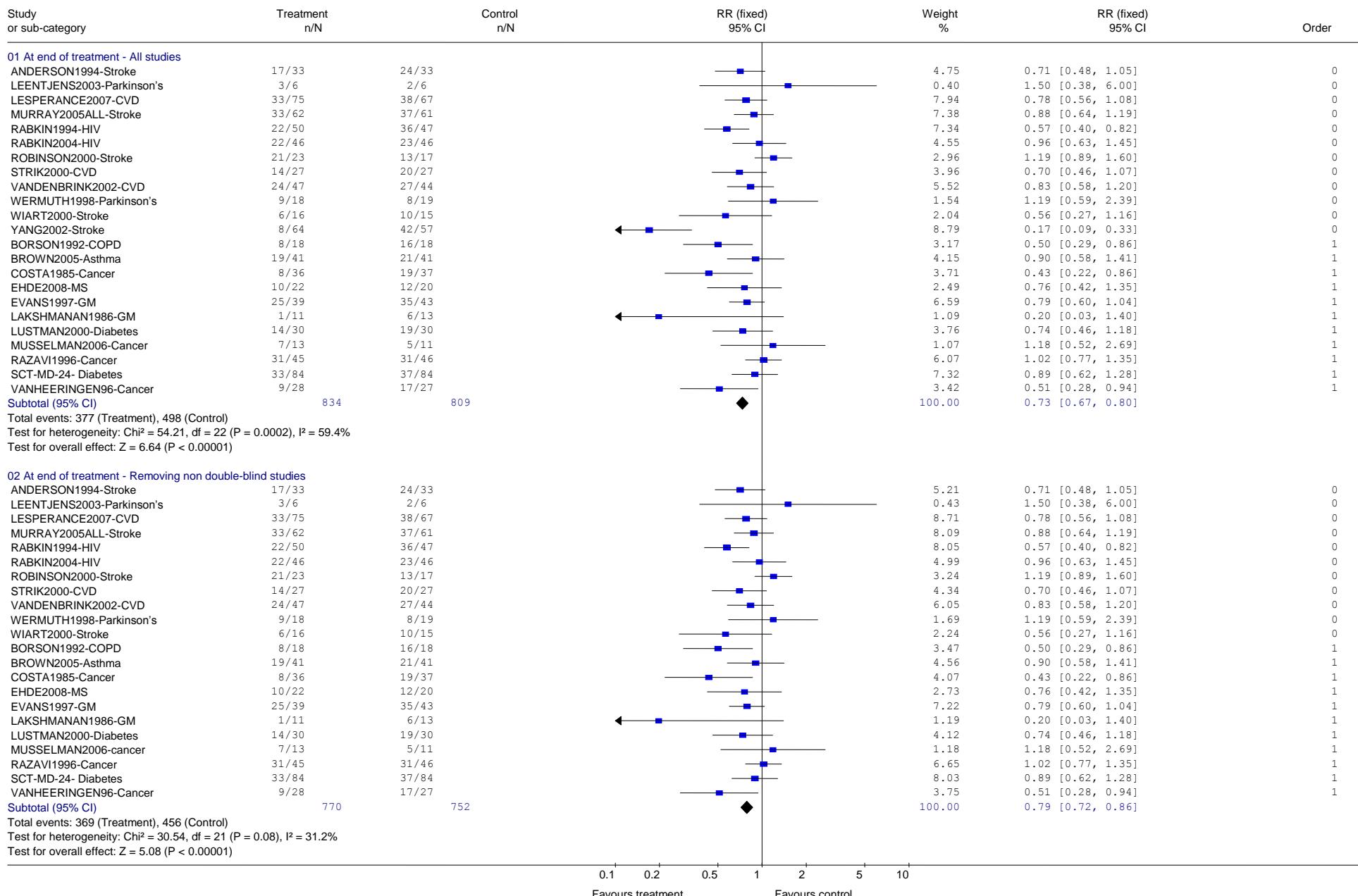
Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 04 Leaving the study early: Due to adverse events



Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 31 Depression: 1. Not achieving success/ remission (reaching a specified cut-off) - observer rated



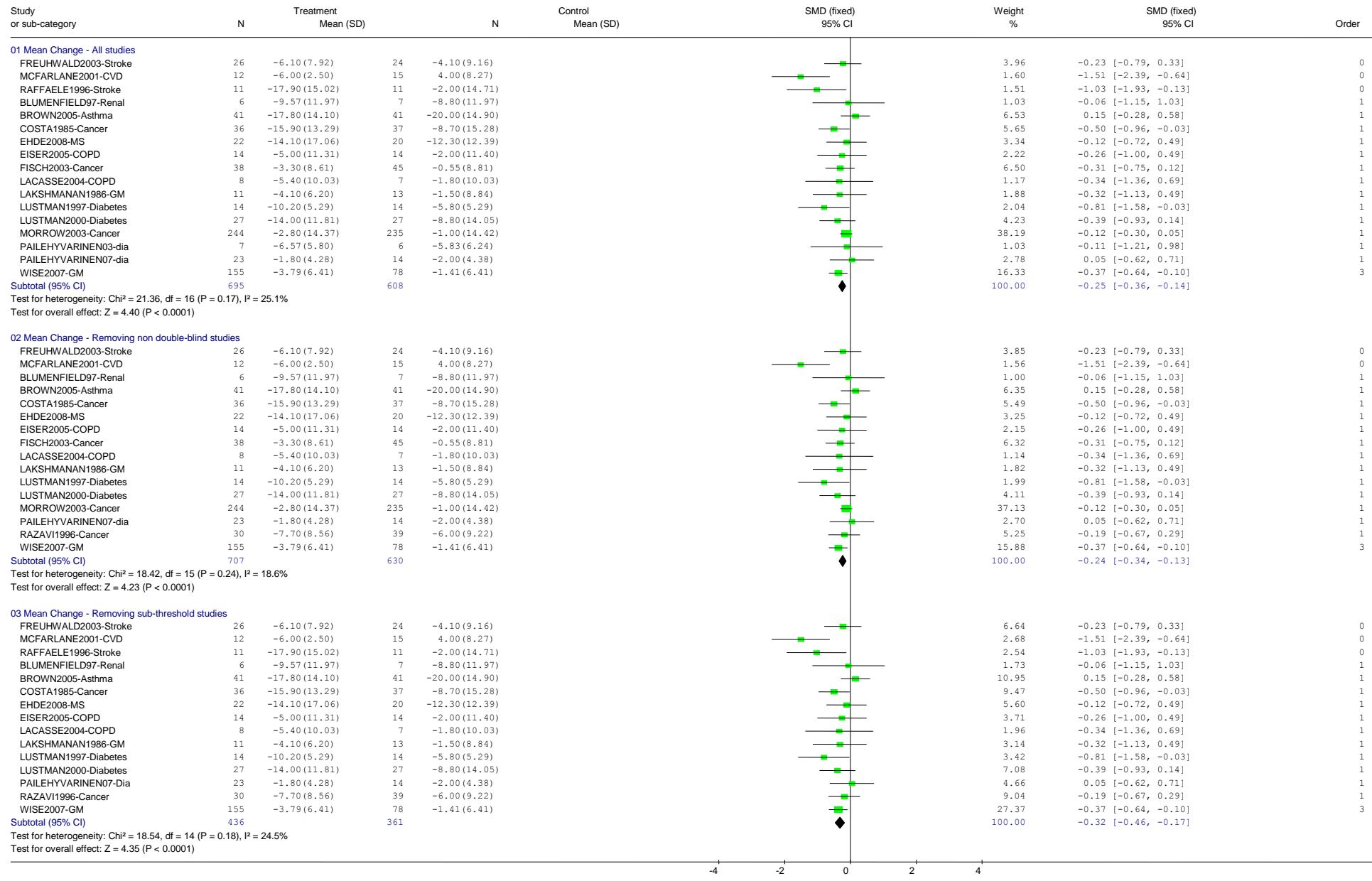
Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 33 Depression: 2. Non-response (not achieving 50% reduction from baseline) - observer rated e.g. HAMD, MADRS



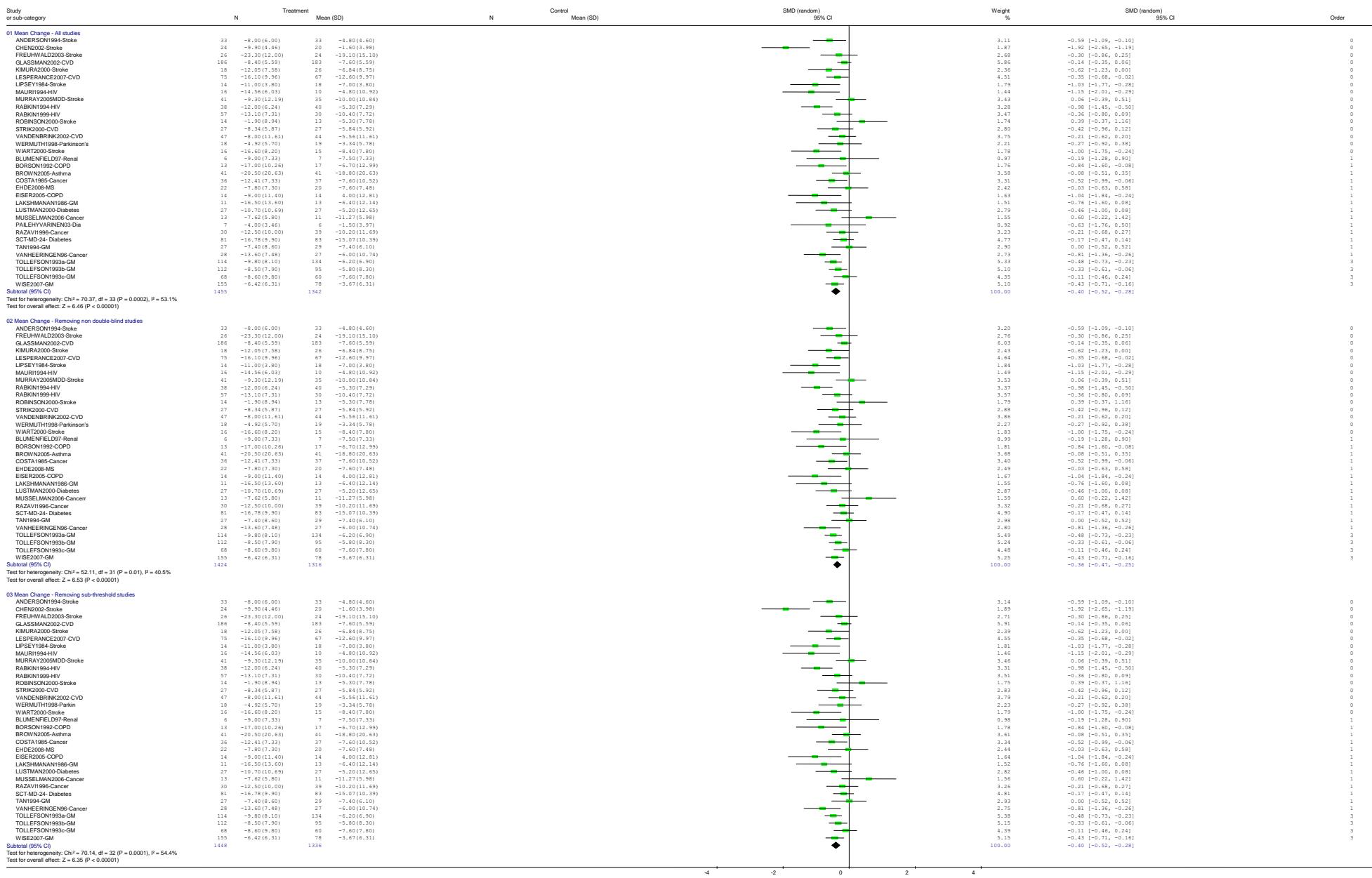
## **Any antidepressant versus placebo – sensitivity analysis removing subthreshold studies**

See next page.

Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 34 Depression: 3. Patient-rated continuous measures

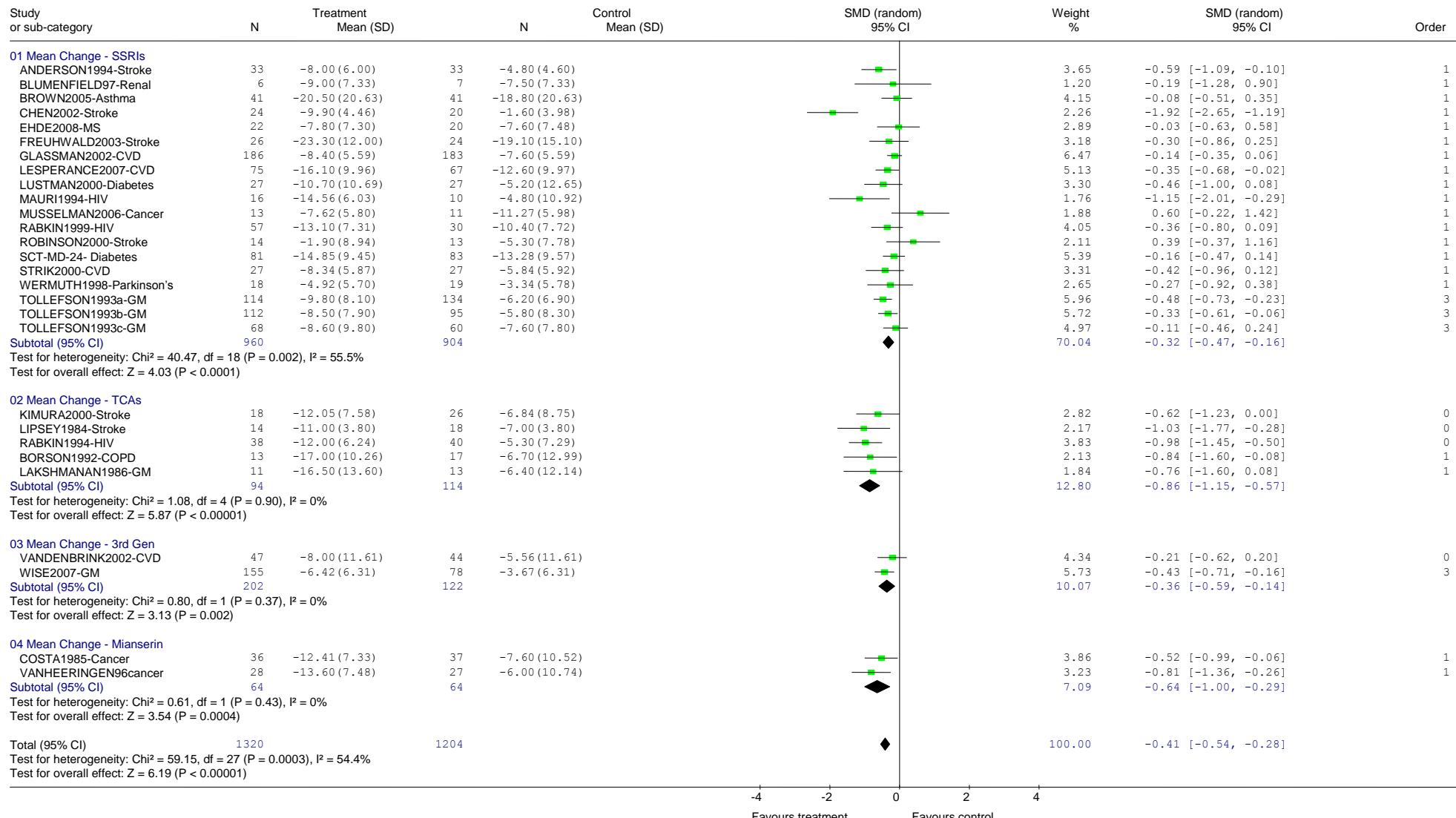


Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 35 Depression: 4. Observer-rated Continuous measures

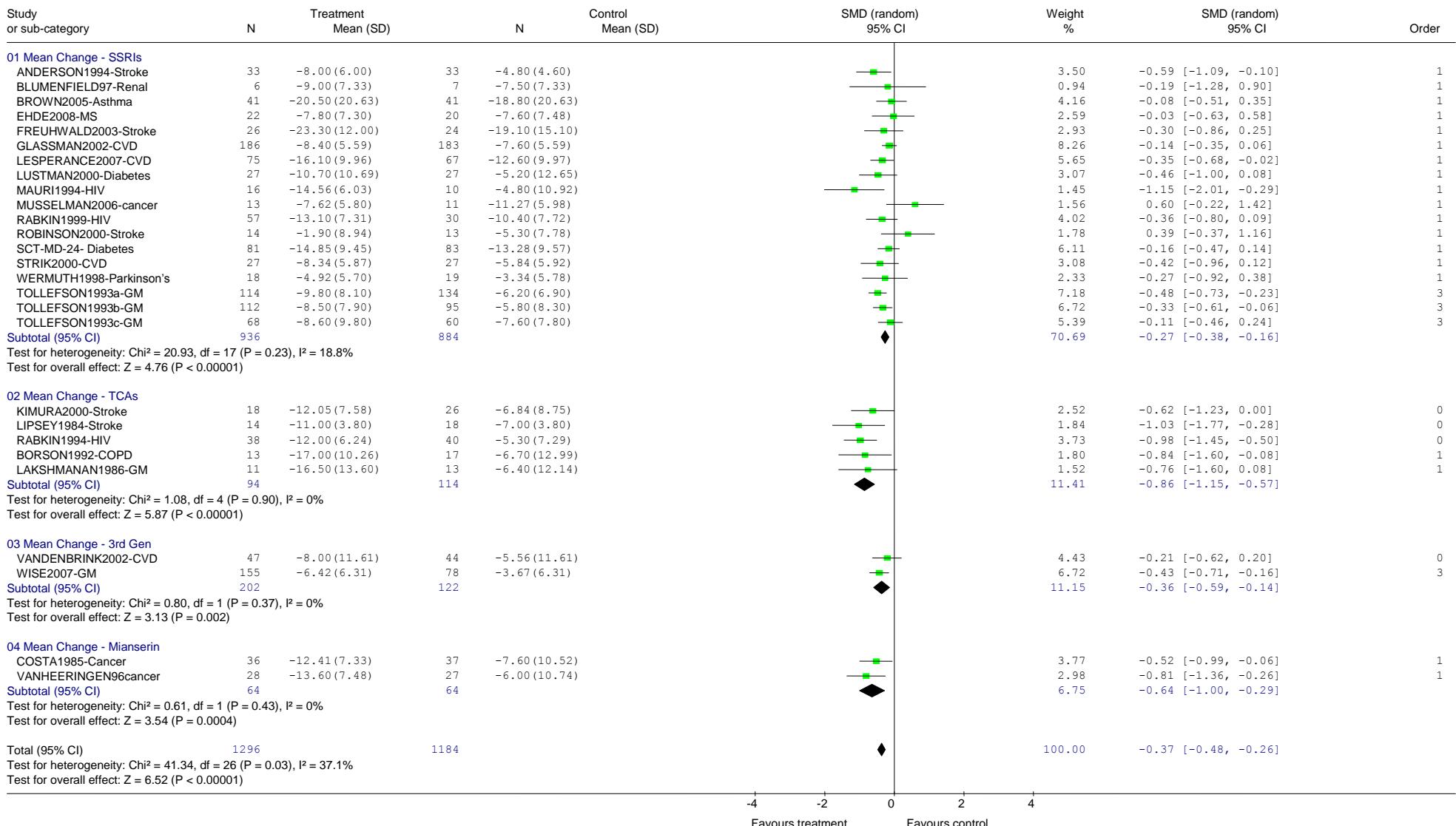


## Any antidepressant versus placebo – subgroup analysis: HAM-D only

Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 36 Depression: 4. Observer-rated continuous measures (HAM-D only)



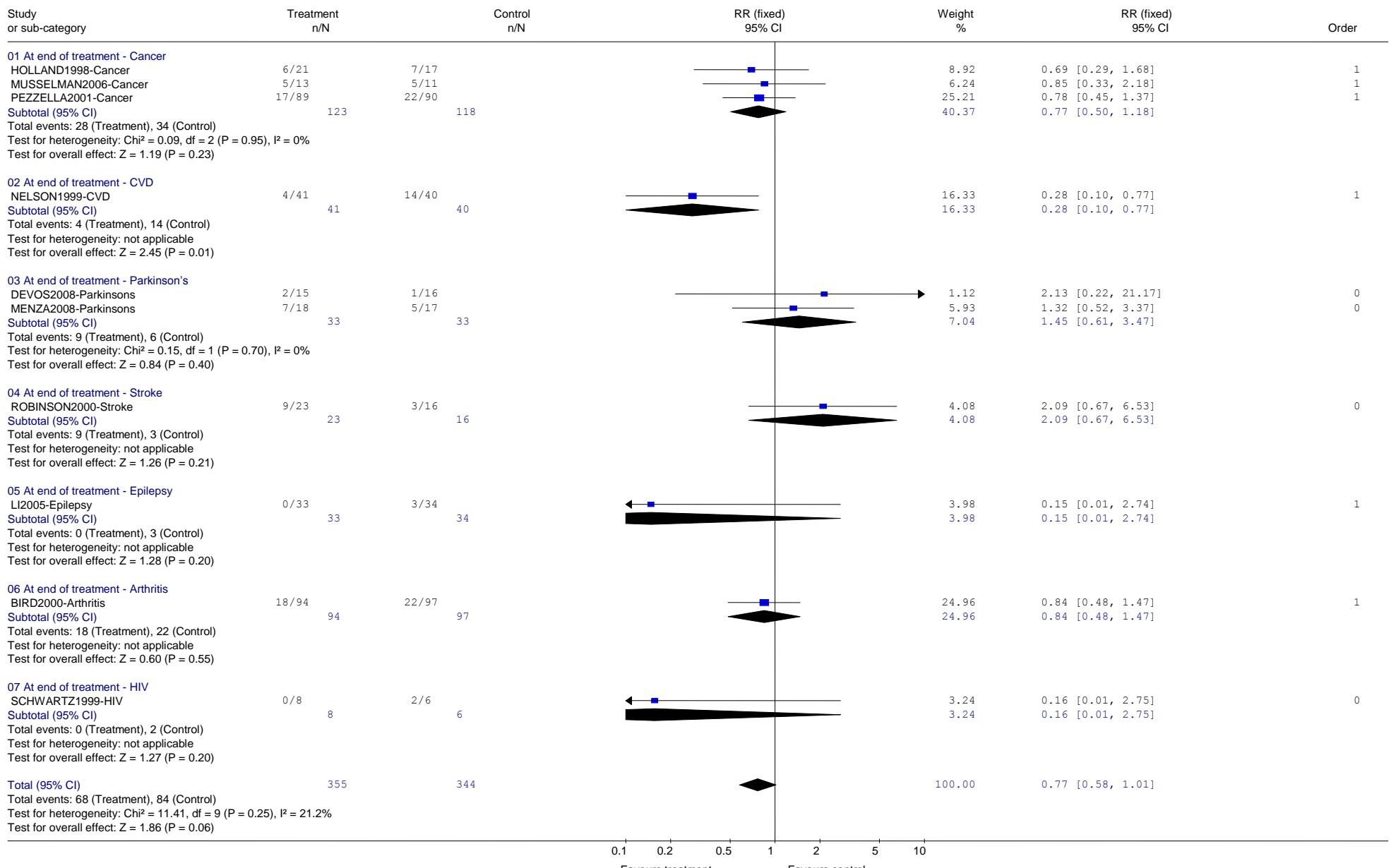
Review: DCHP - Pharm - by drug class  
 Comparison: 11 Any antidepressant versus placebo (sensitivity analyses)  
 Outcome: 37 Depression: 4. Observer-rated continuous measures (HAM-D only - removing non double-blind and sub-threshold)



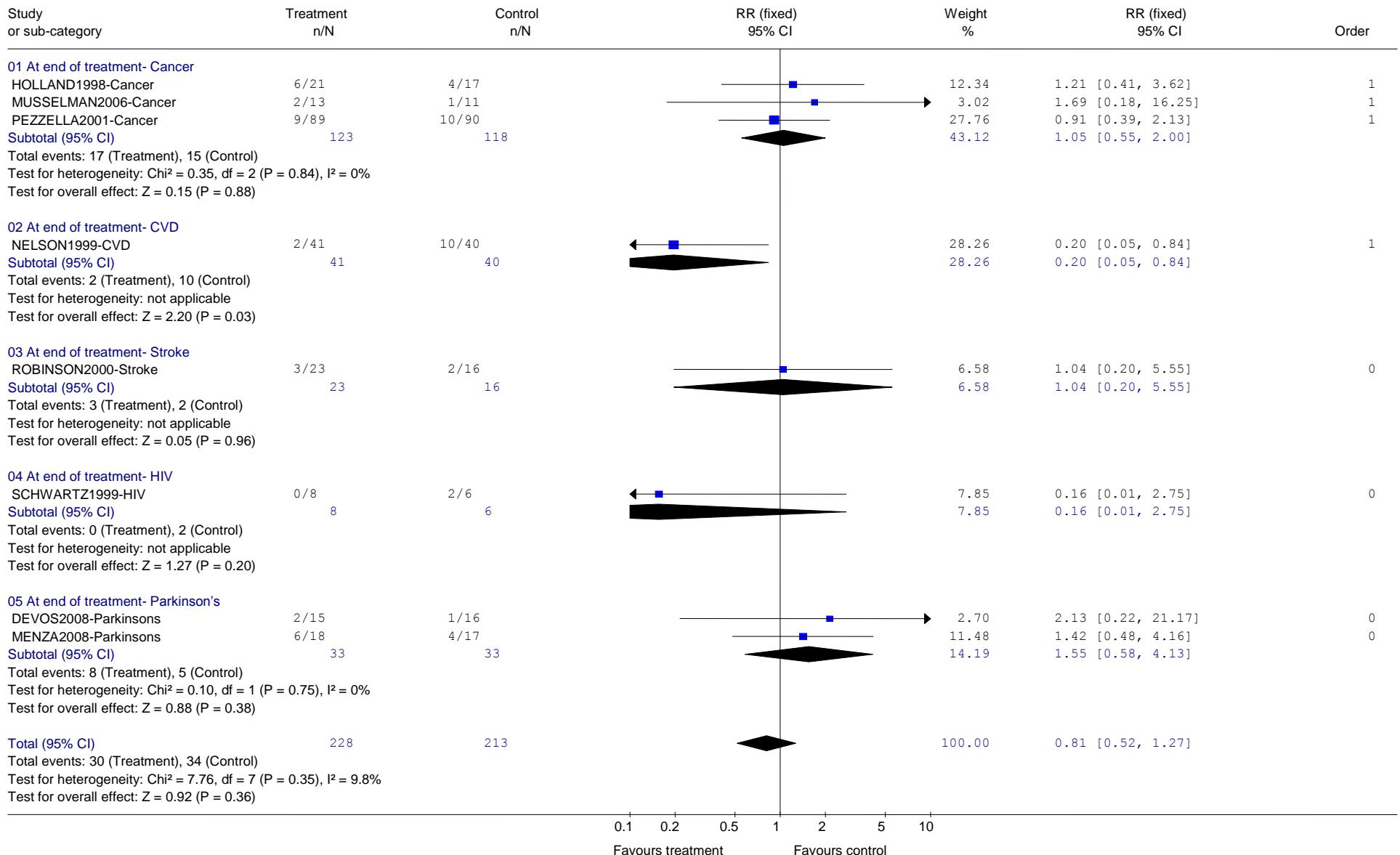
## **SSRIs versus TCAs**

See next page.

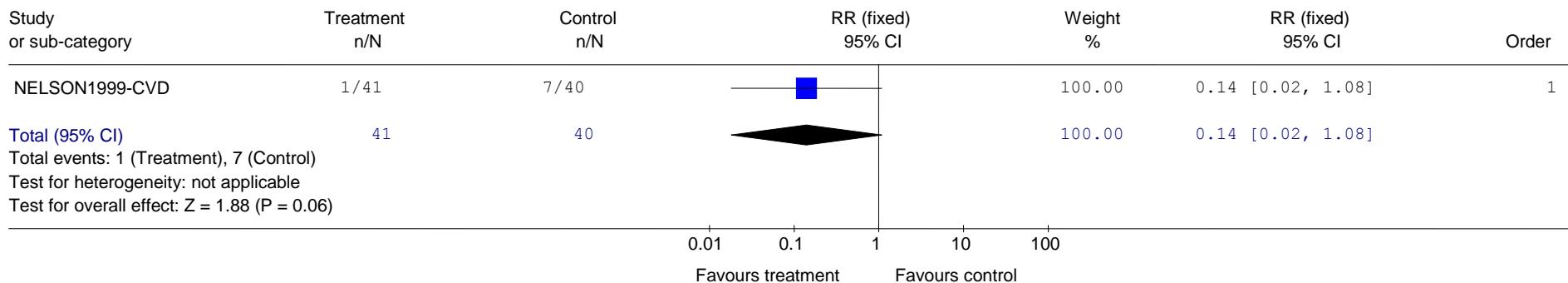
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 20 Leaving the study early - any reason



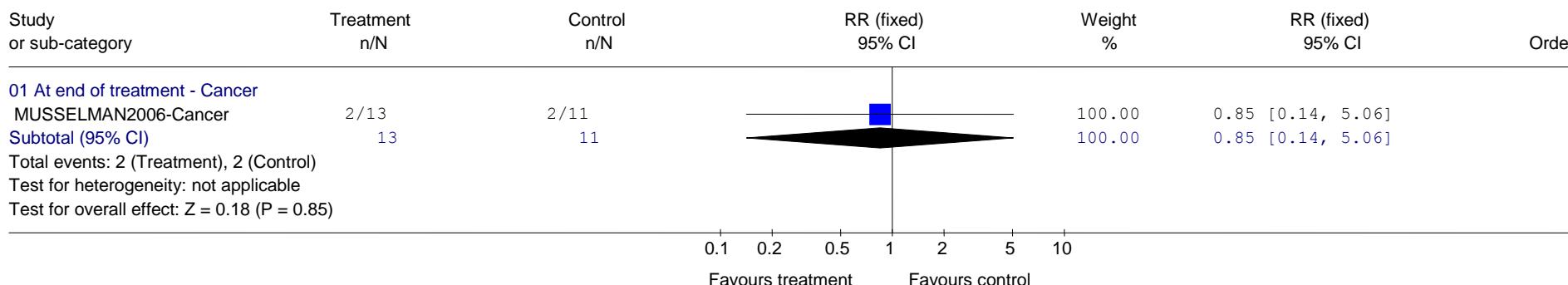
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 21 Leaving study early due to adverse events



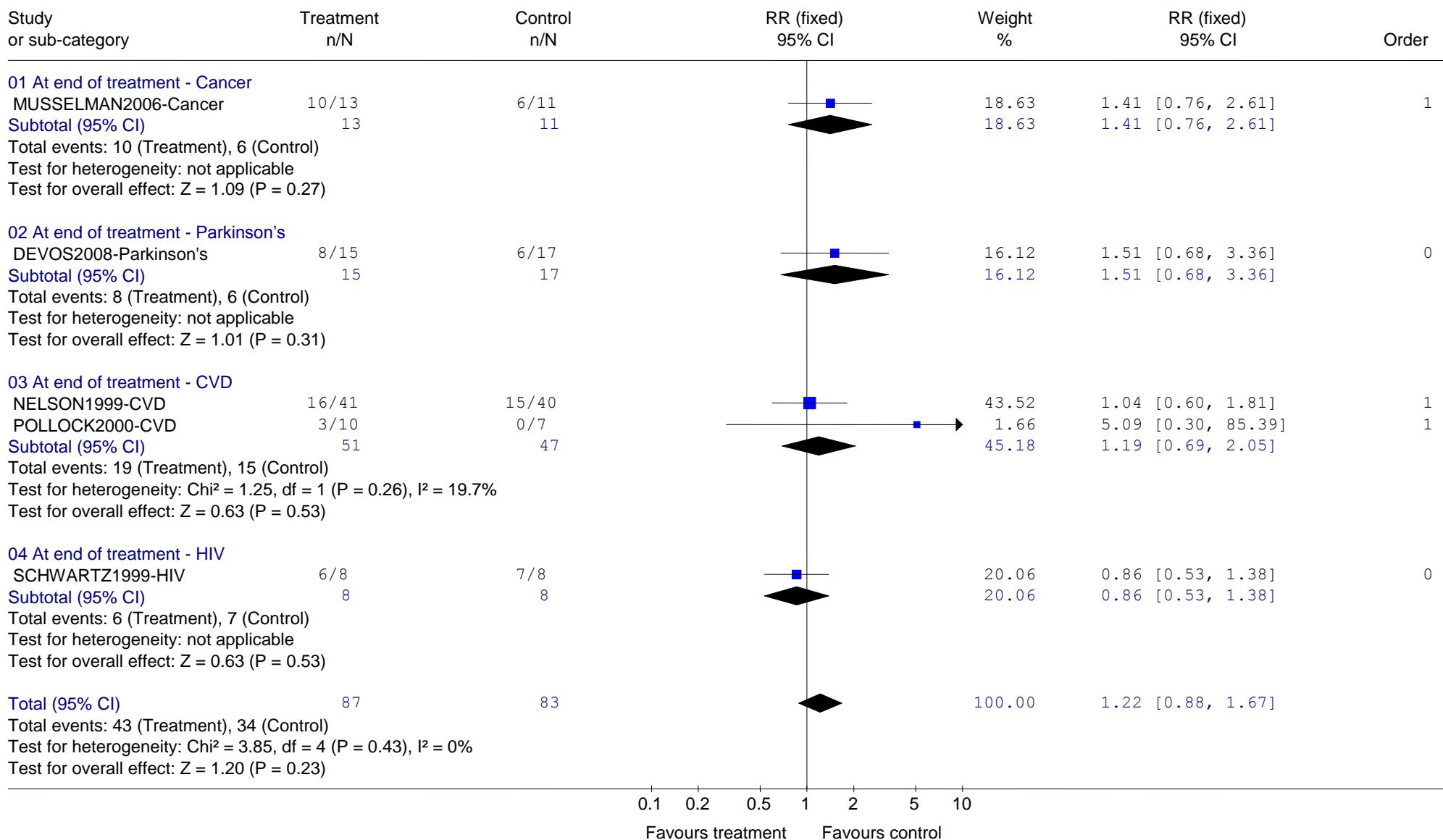
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 22 Leaving study early due to adverse cardiac events



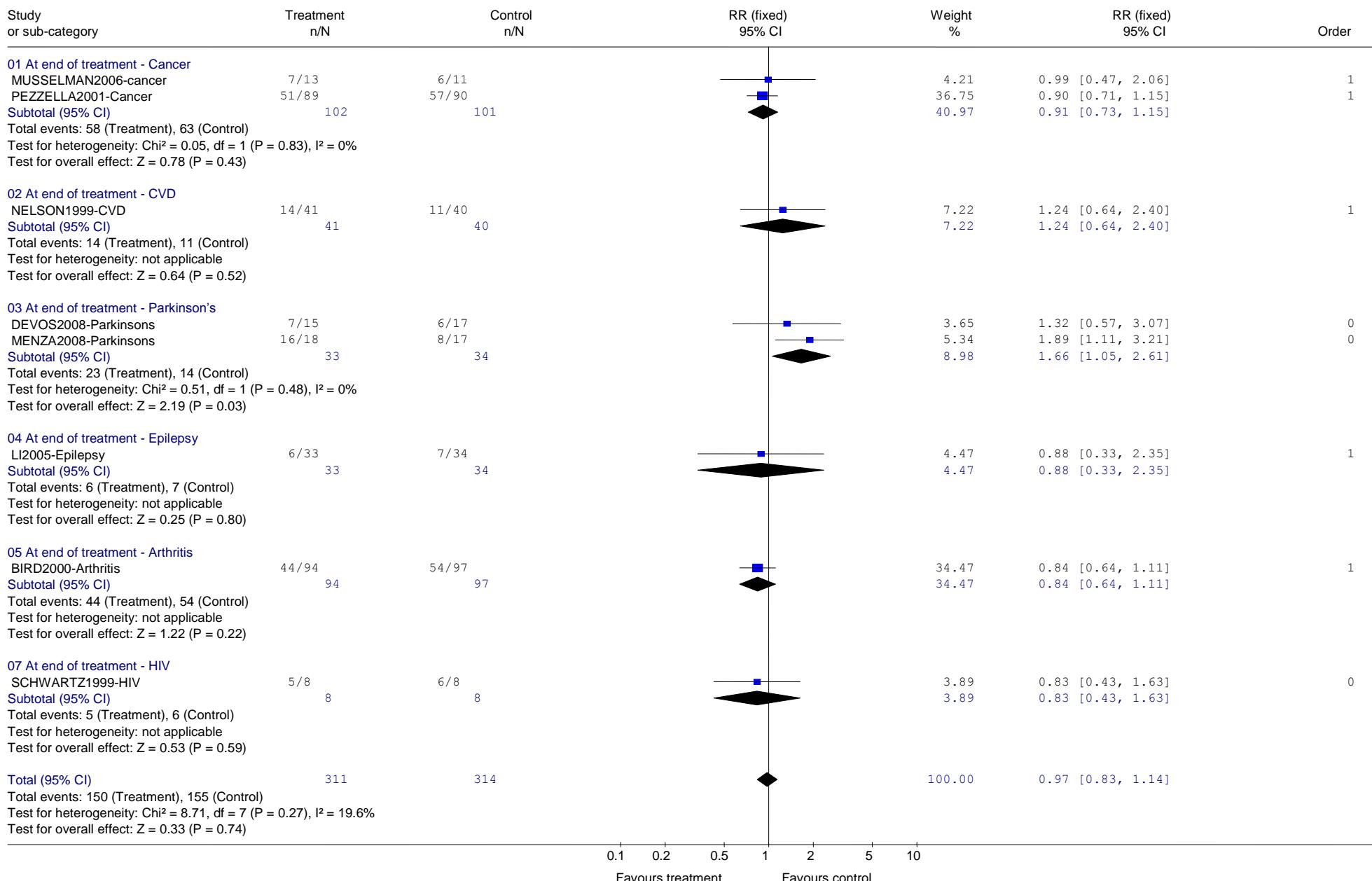
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 23 Leaving the study early: Due to lack of efficacy



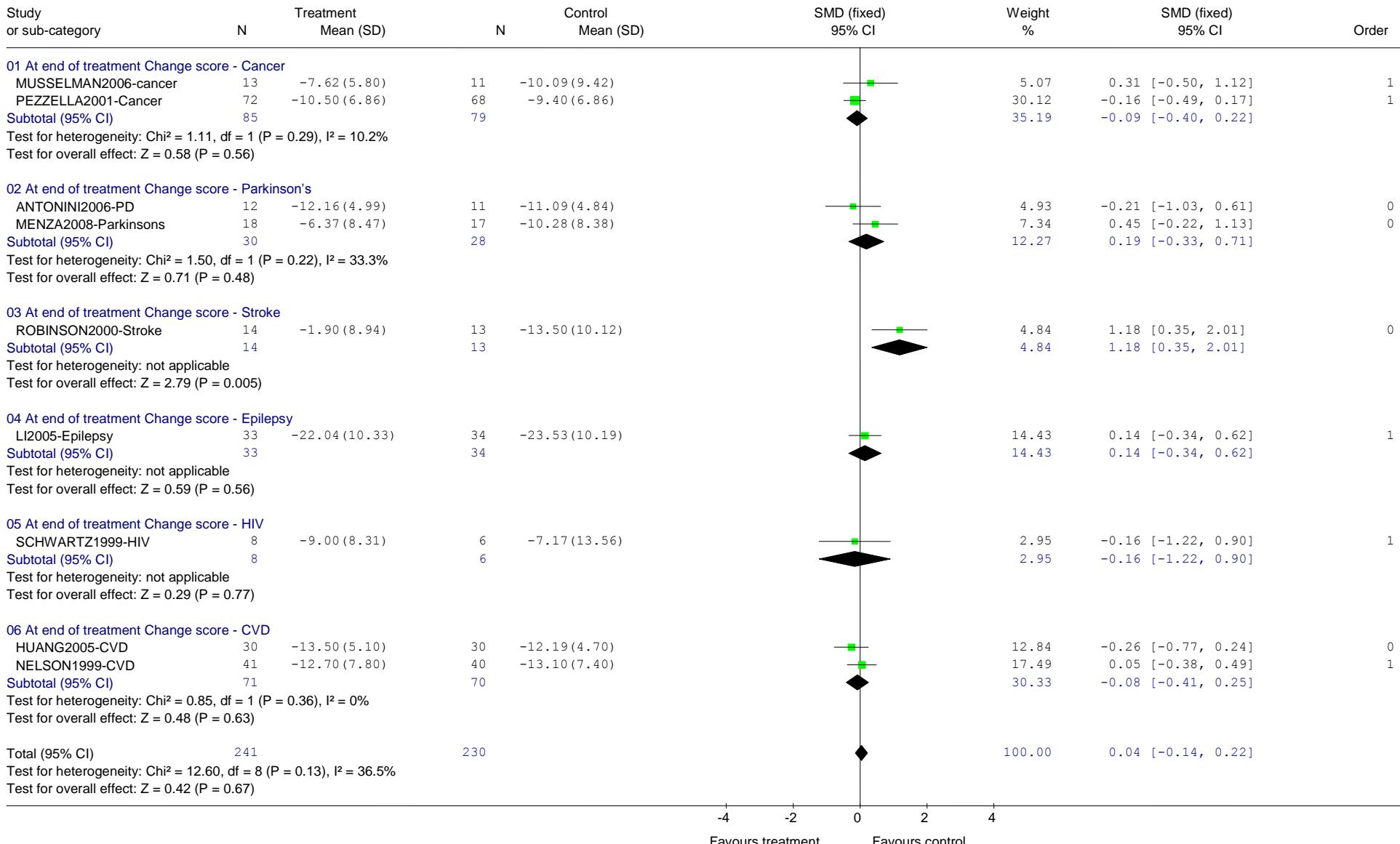
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 30 Depression: 1. Remission (below cut-off)



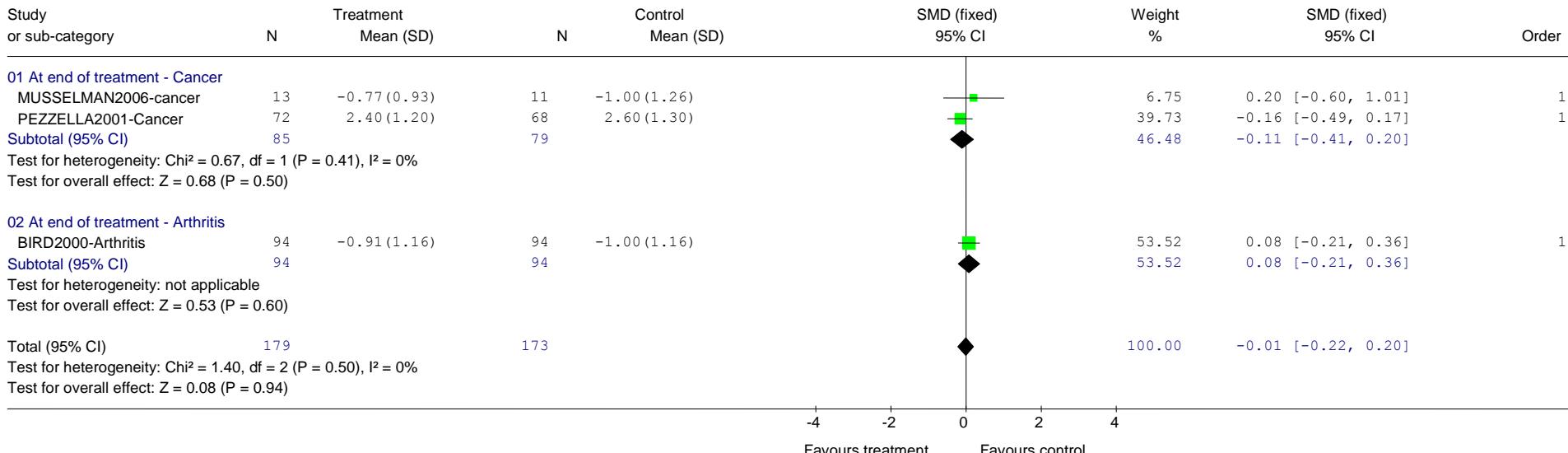
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 31 Depression: 2. Non-response (<50% reduction)



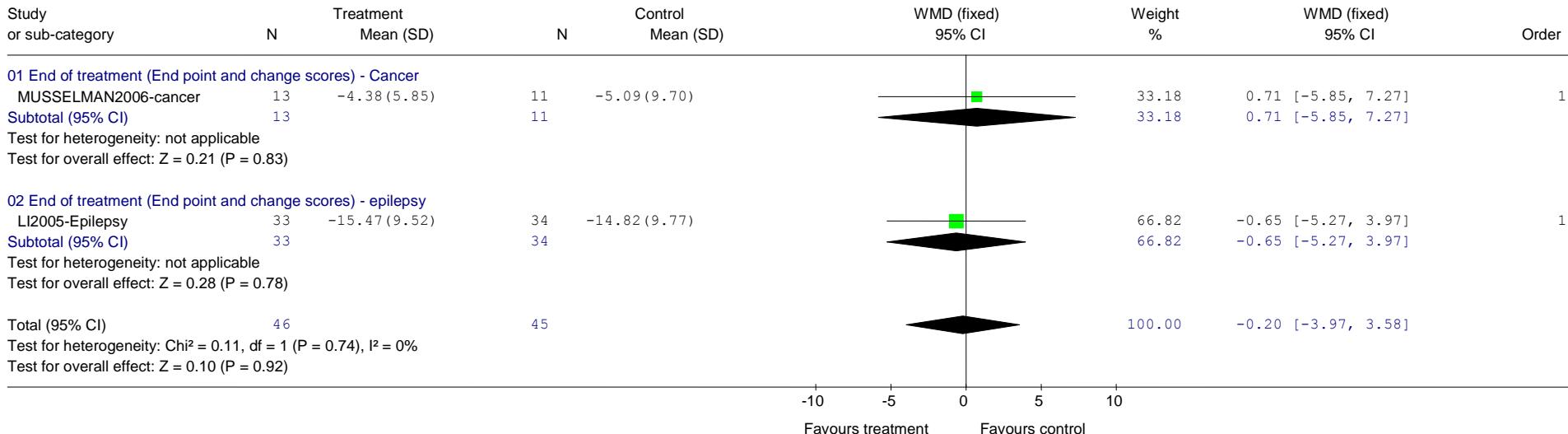
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 32 Depression: 3. Continuous measures



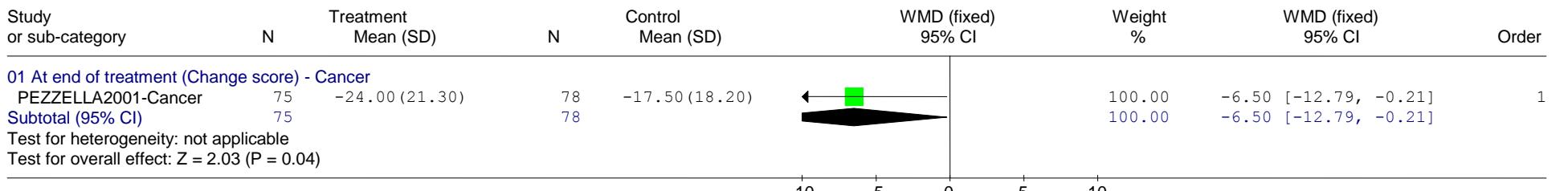
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 40 Global state: 1. CGI



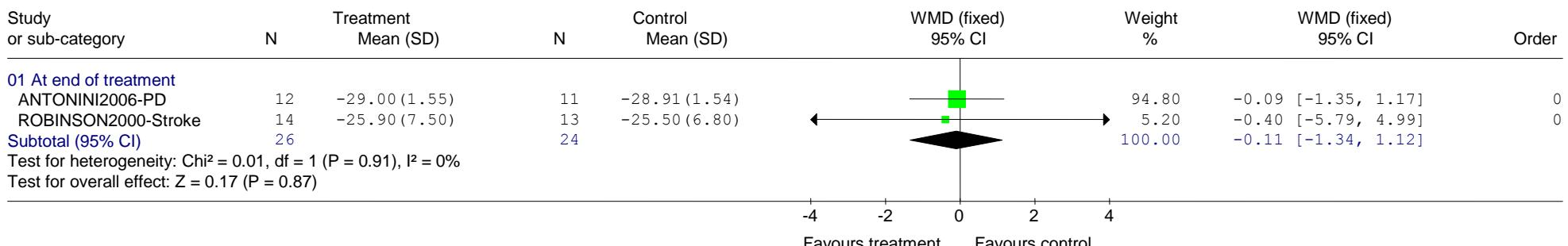
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 41 Mental State: 1. Continuous measures - HAM-A



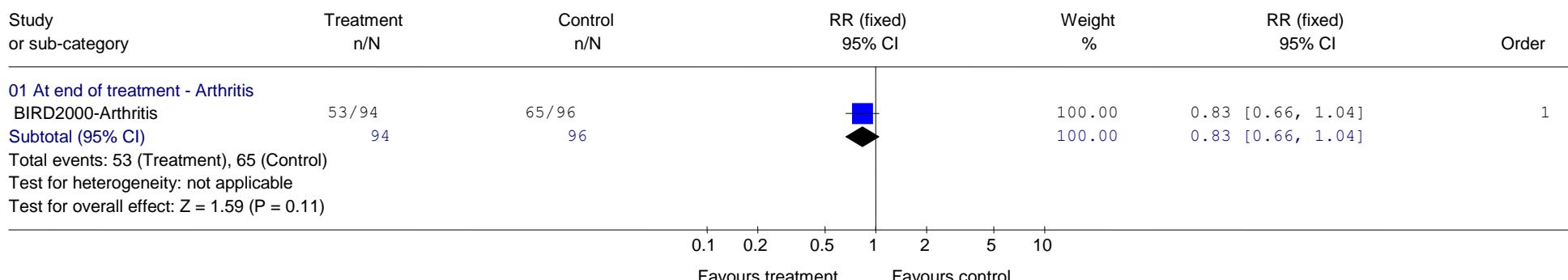
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 50 QoL: 1. Continuous measures



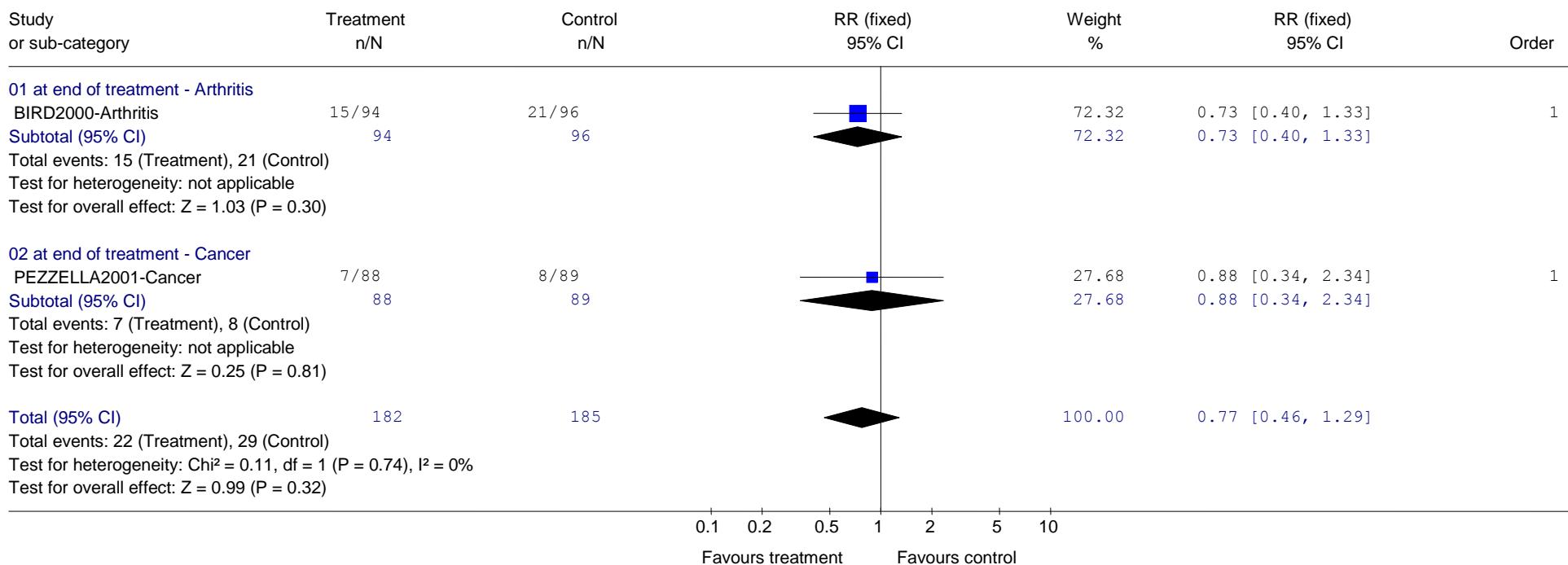
Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 55 Physical health outcome: 1. MMSE - PD



Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 60 Adverse events - reporting at least 1 adverse event

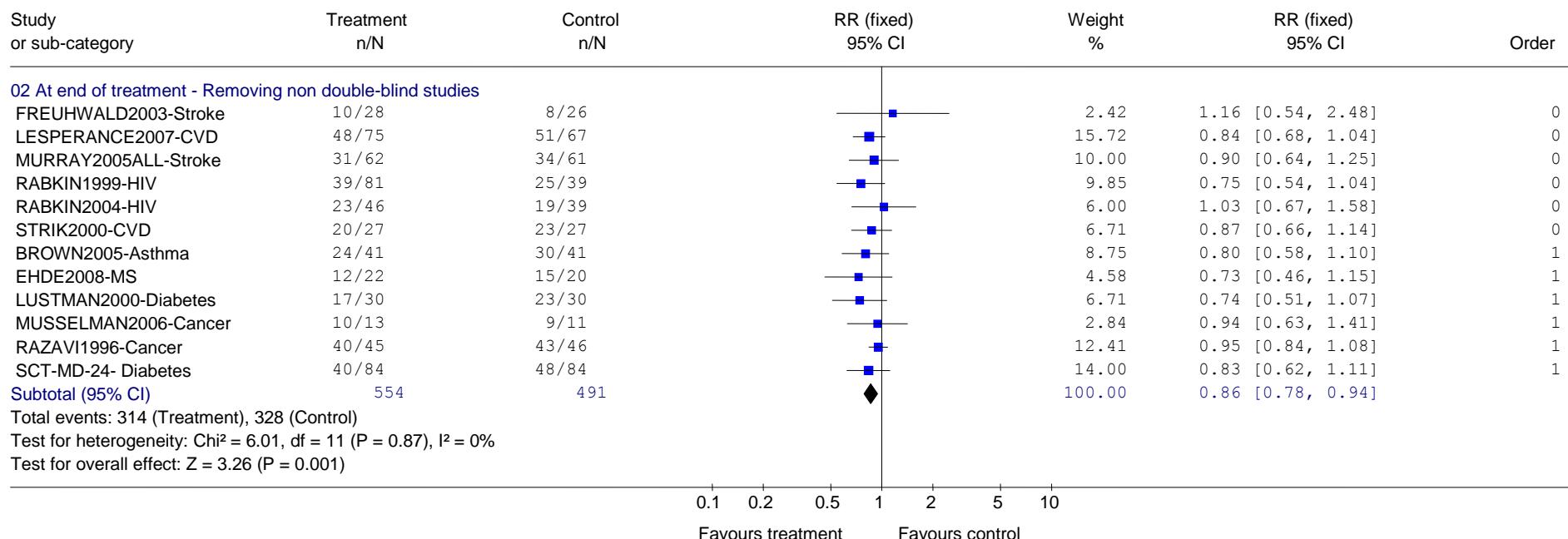


Review: DCHP - Pharm - by drug class  
 Comparison: 20 Head-to-head SSRI versus TCA  
 Outcome: 61 Adverse events - severe adverse event

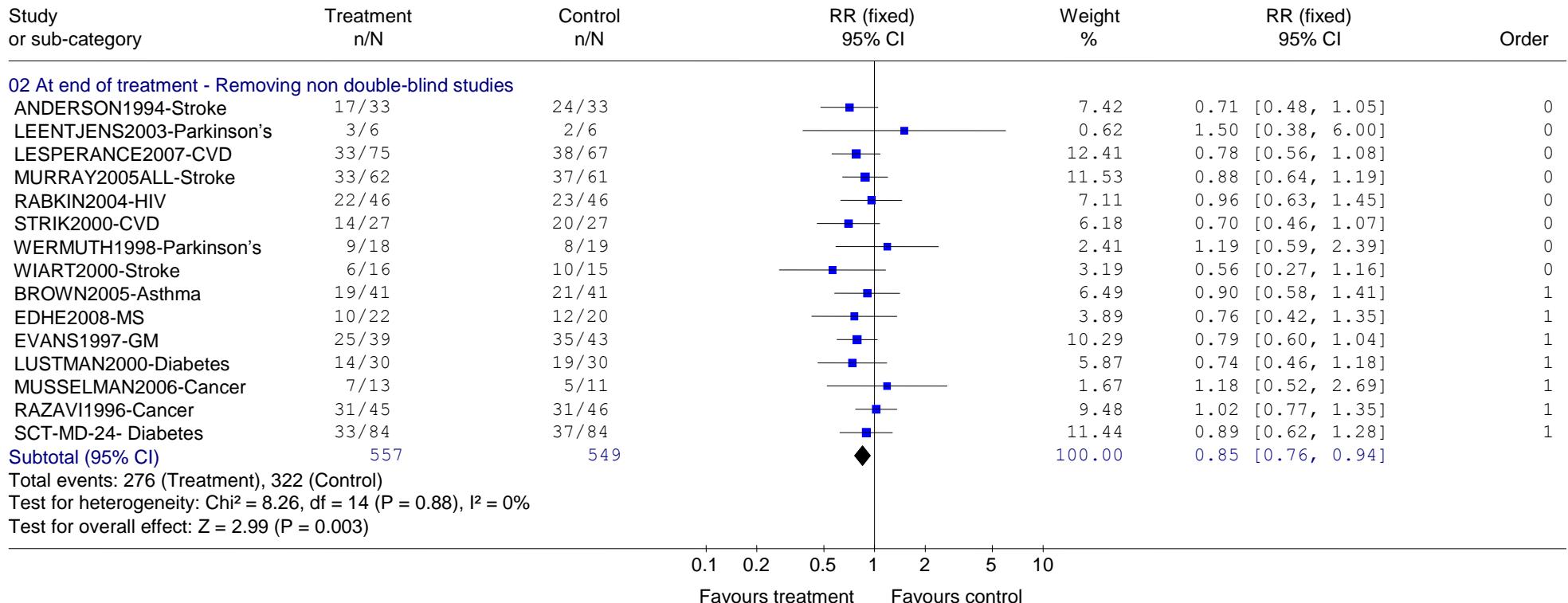


## SSRI versus placebo - sensitivity analysis removing non-blinded and subthreshold studies

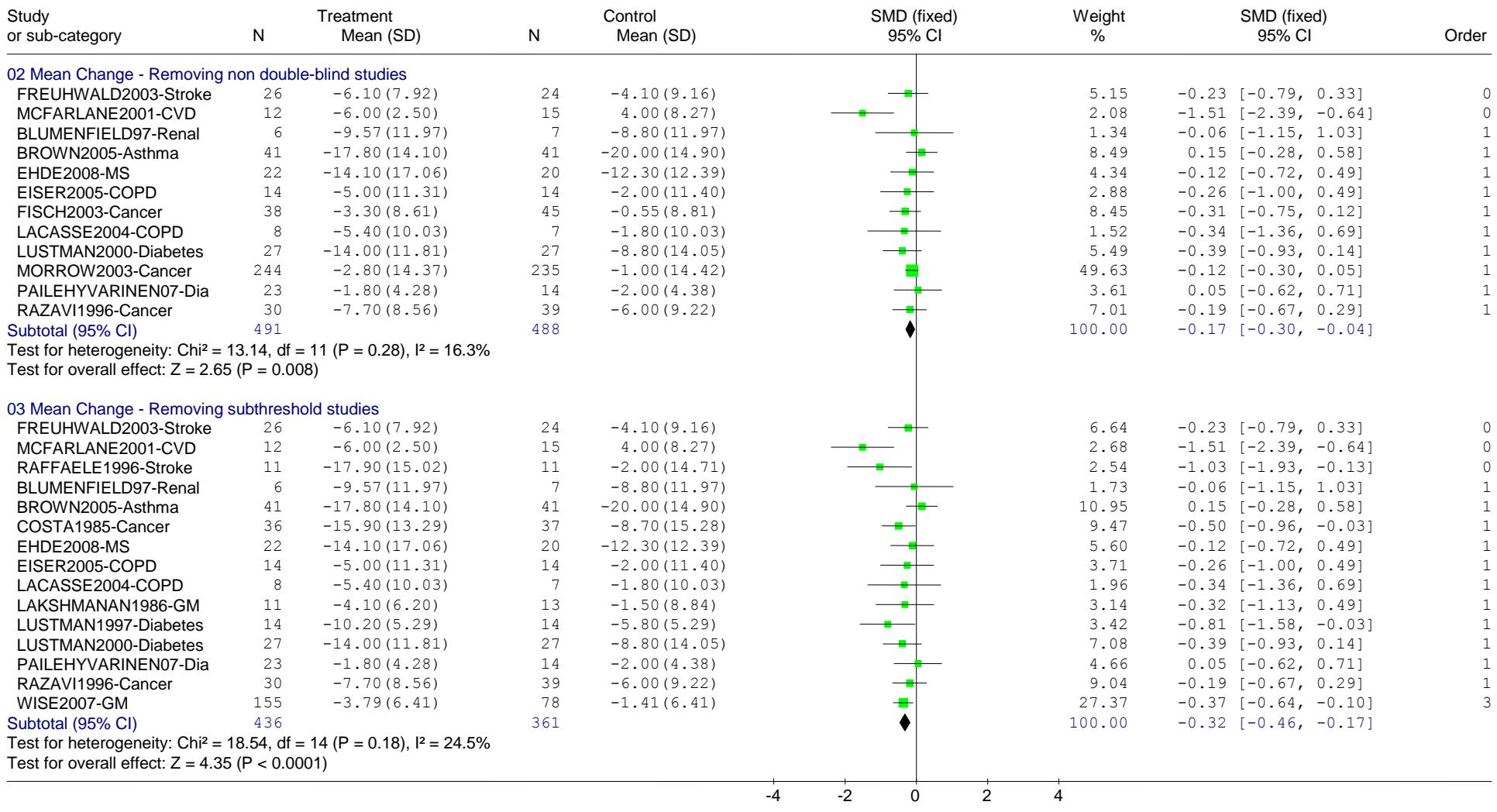
Review: DCHP - Pharm - by drug class  
 Comparison: 32 SSRIs versus placebo (Sensitivity analyses: Blinded/Non-blinded)  
 Outcome: 01 Depression: 1. Not achieving success/ remission (reaching a specified cut-off) - observer rated



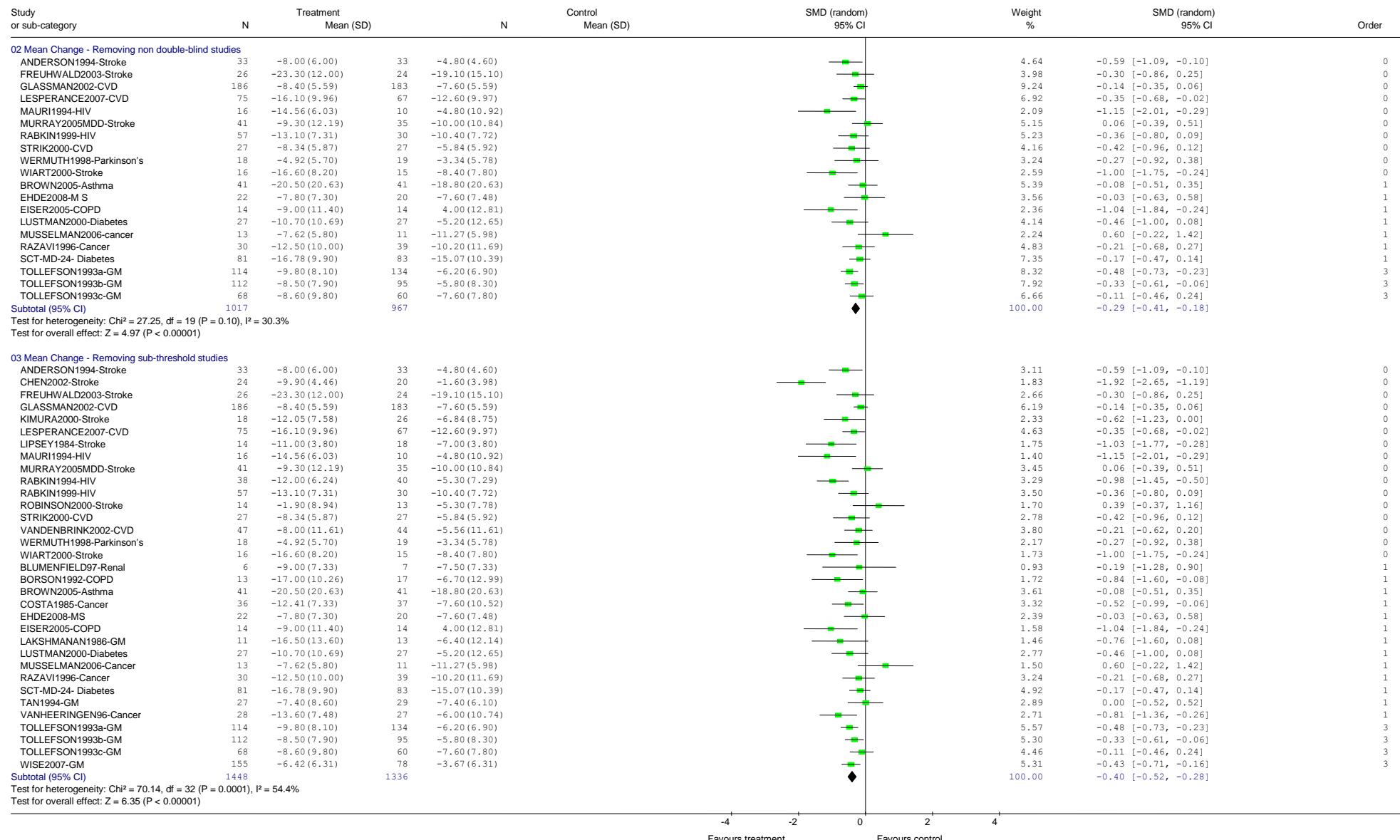
Review: DCHP - Pharm - by drug class  
 Comparison: 32 SSRIs versus placebo (Sensitivity analyses: Blinded/Non-blinded)  
 Outcome: 02 Depression: 2. Non-response (not achieving 50% reduction from baseline) - observer rated e.g. HAM-D, MADRS



Review: DCHP - Pharm - by drug class  
 Comparison: 32 SSRIs versus placebo (Sensitivity analyses: Blinded/Non-blinded)  
 Outcome: 03 Depression: 3. Patient-rated Continuous measures



Review: DCHP - Pharm - by drug class  
 Comparison: 32 SSRIs versus placebo (Sensitivity analyses: Blinded/Non-blinded)  
 Outcome: 04 Depression: 4. Observer-rated continuous measures

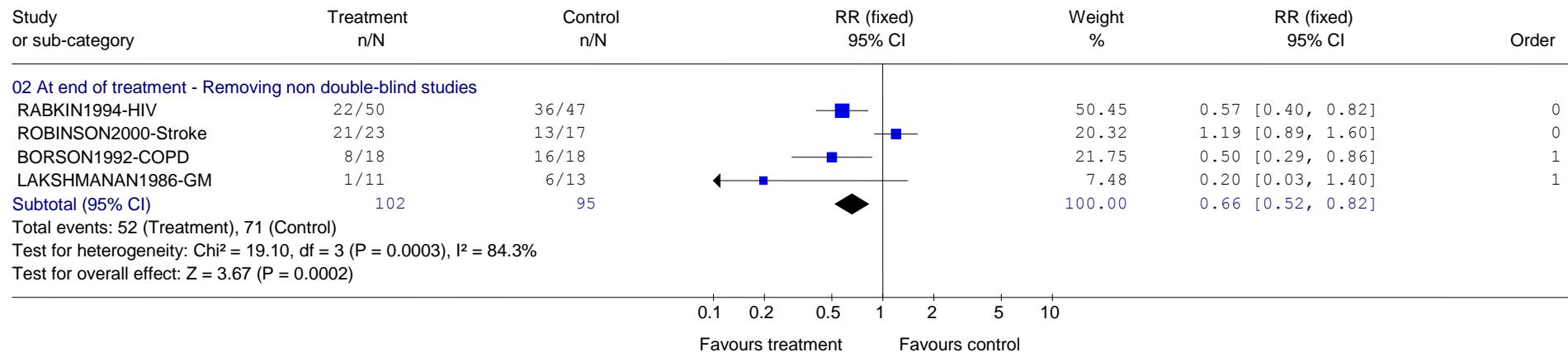


## TCAs versus placebo - sensitivity analysis removing non-blinded and subthreshold studies

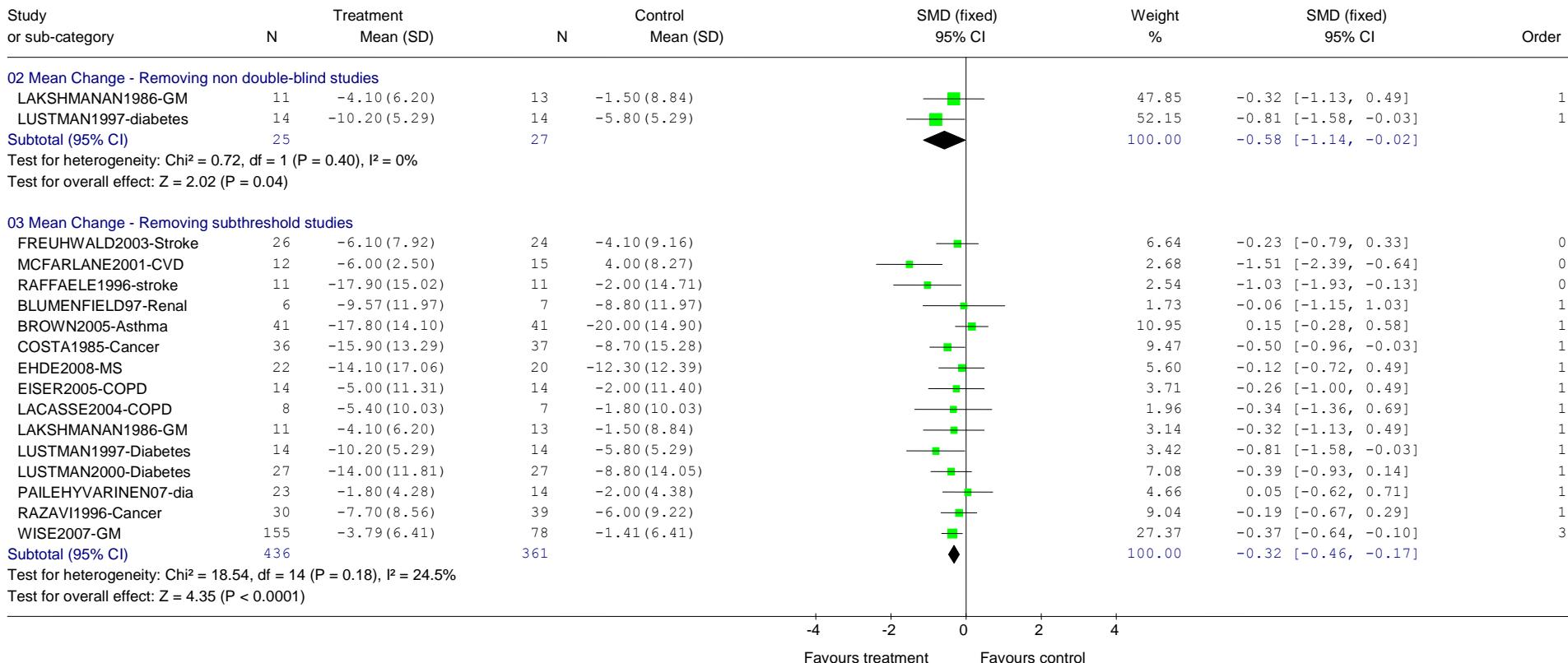
Review: DCHP - Pharm - by drug class

Comparison: 33 TCAs versus placebo (Sensitivity analyses: Blinded/Non-blinded)

Outcome: 02 Depression: 2. Non-response (not achieving 50% reduction from baseline) - observer rated e.g. HAMD, MADRS



Review: DCHP - Pharm - by drug class  
 Comparison: 33 TCAs versus Placebo (Sensitivity analyses: Blinded/non-blinded)  
 Outcome: 03 Depression: 3. Patient-rated continuous measures



Review: DCHP - Pharm - by drug class  
 Comparison: 33 TCAs versus placebo (Sensitivity analyses: Blinded/Non-blinded)  
 Outcome: 04 Depression: 4. Observer-rated Continuous measures

