

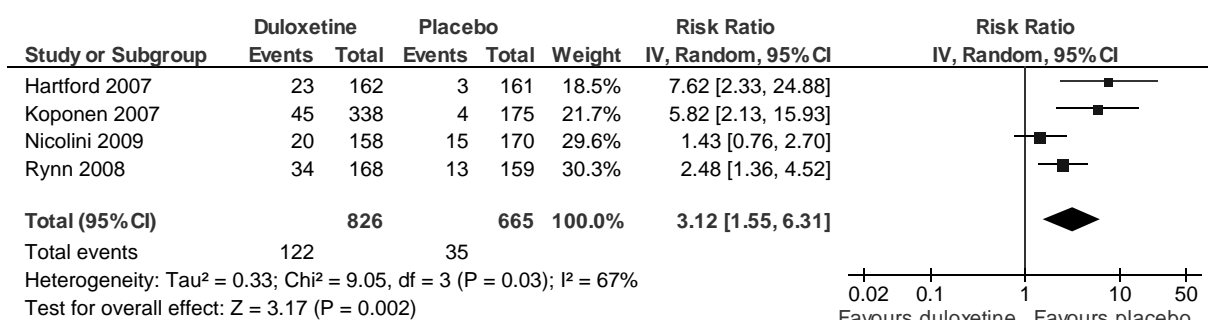
## Appendix 16c: Pharmacological interventions forest plots

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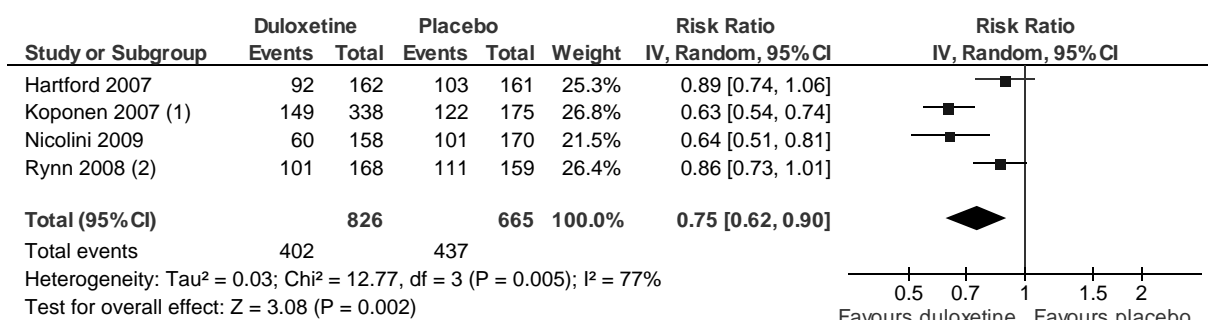
# 1 Antidepressants versus Placebo for GAD

## 1.1 Duloxetine versus Placebo

### 1.2 Discontinuation due to adverse events



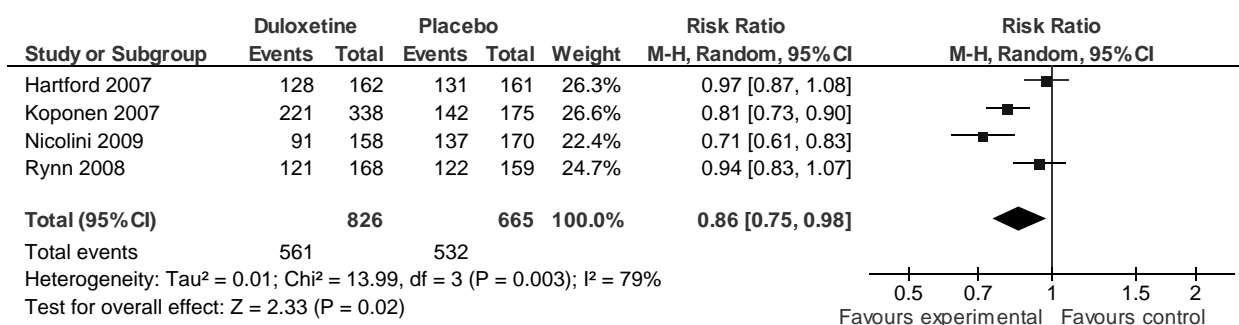
### 1.3 Non-Response



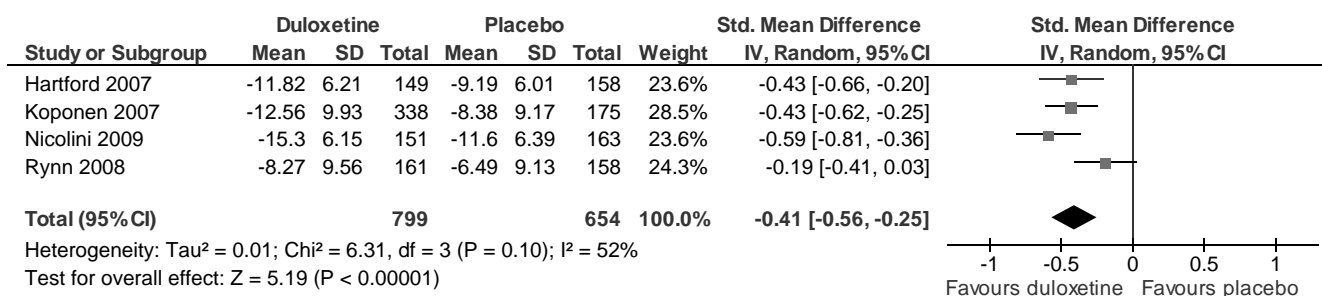
(1) changed

(2) changed

### 1.4 Non-remission

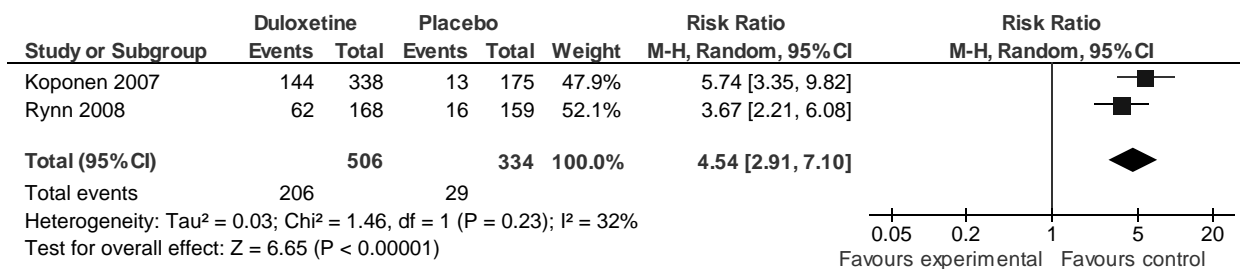


### 1.5 HAM-A Mean change from baseline

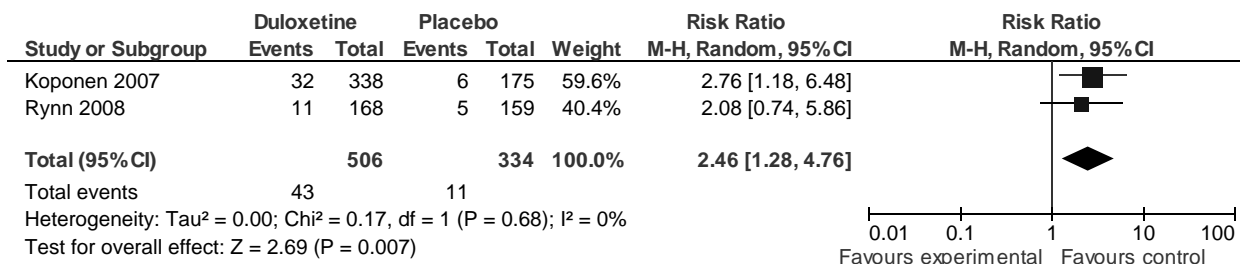


# 1 Antidepressants versus Placebo for GAD

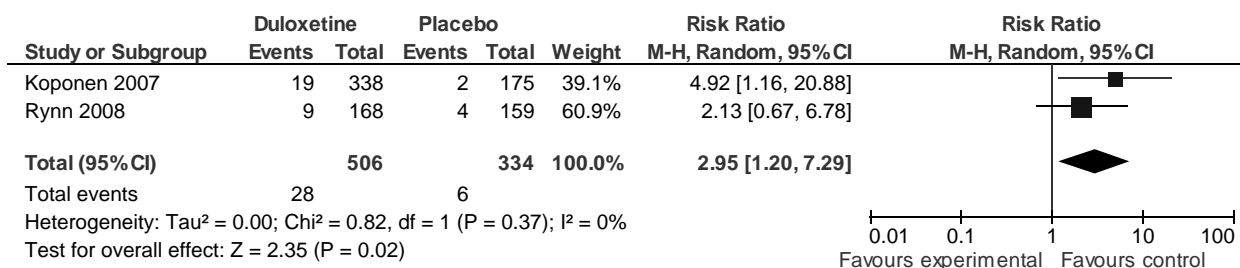
## 1.11 Nausea



## 1.12 Insomnia

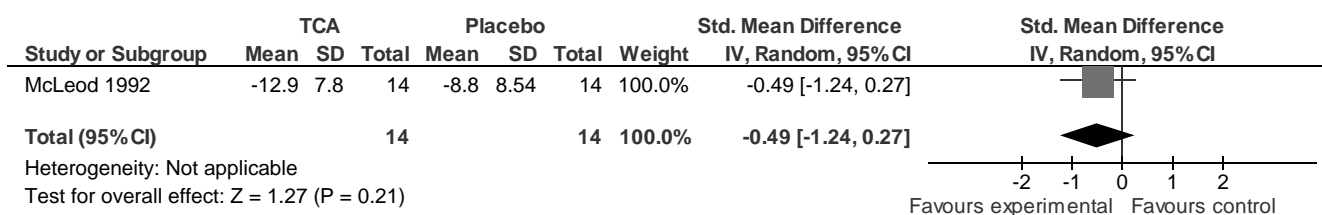


## 1.13 Sexual problems



## 2 Imipramine versus Placebo

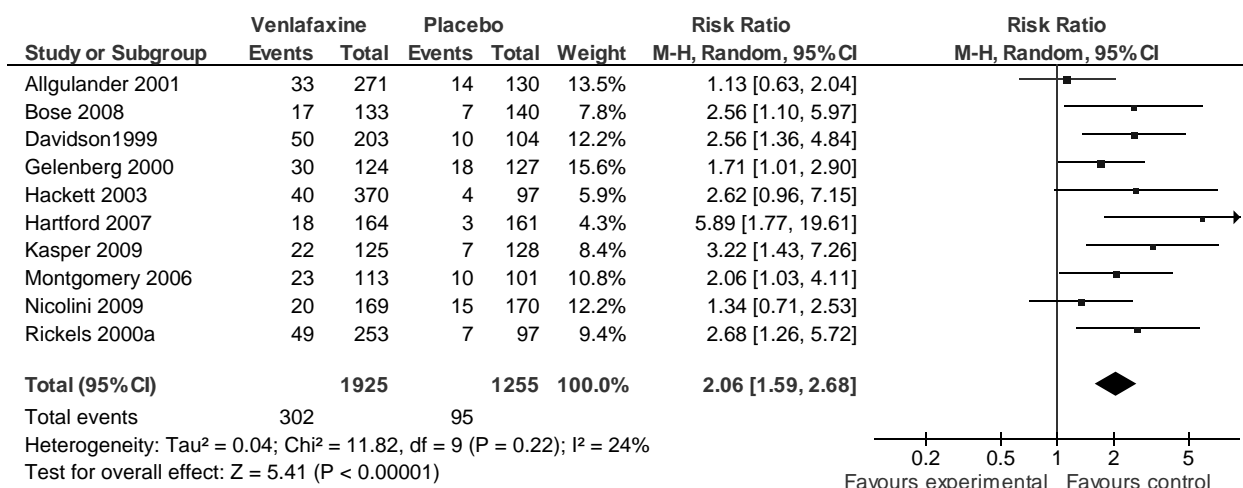
### 2.1 HAM-A



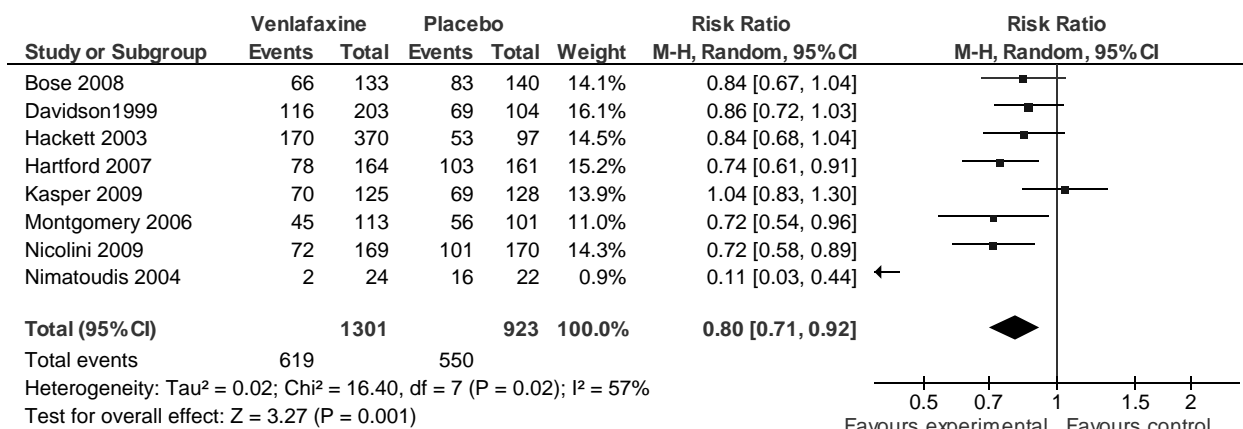
## 3 Venlafaxine versus Placebo

# 1 Antidepressants versus Placebo for GAD

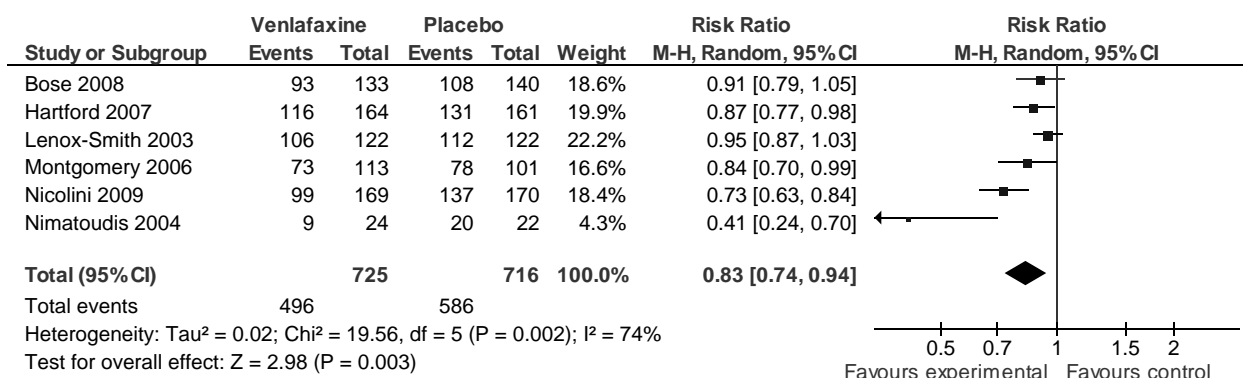
## 3.2 Discontinuation due to adverse events



## 3.3 Non-response

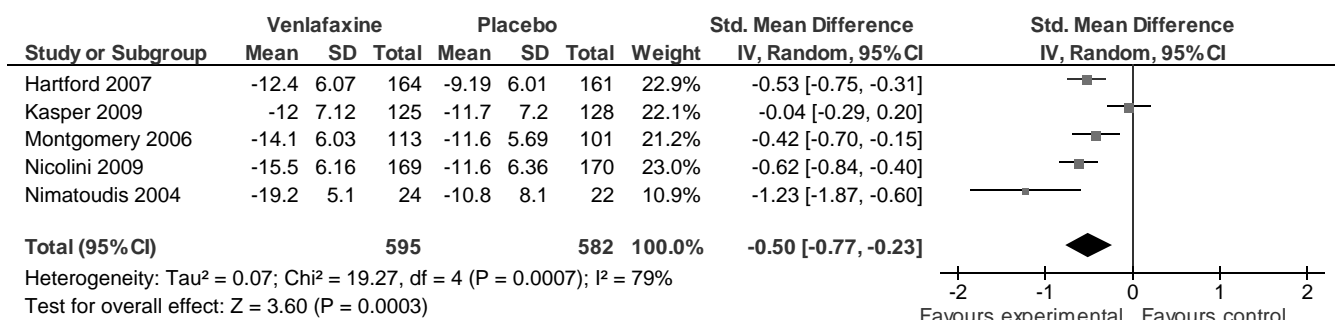


## 3.5 Non-remission

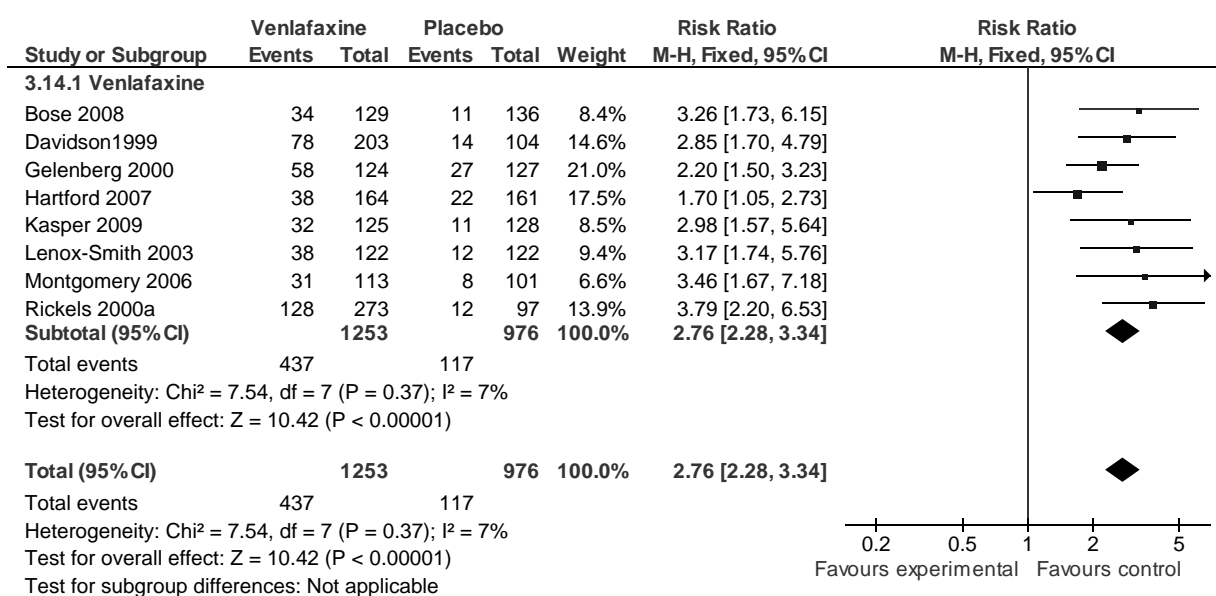


# 1 Antidepressants versus Placebo for GAD

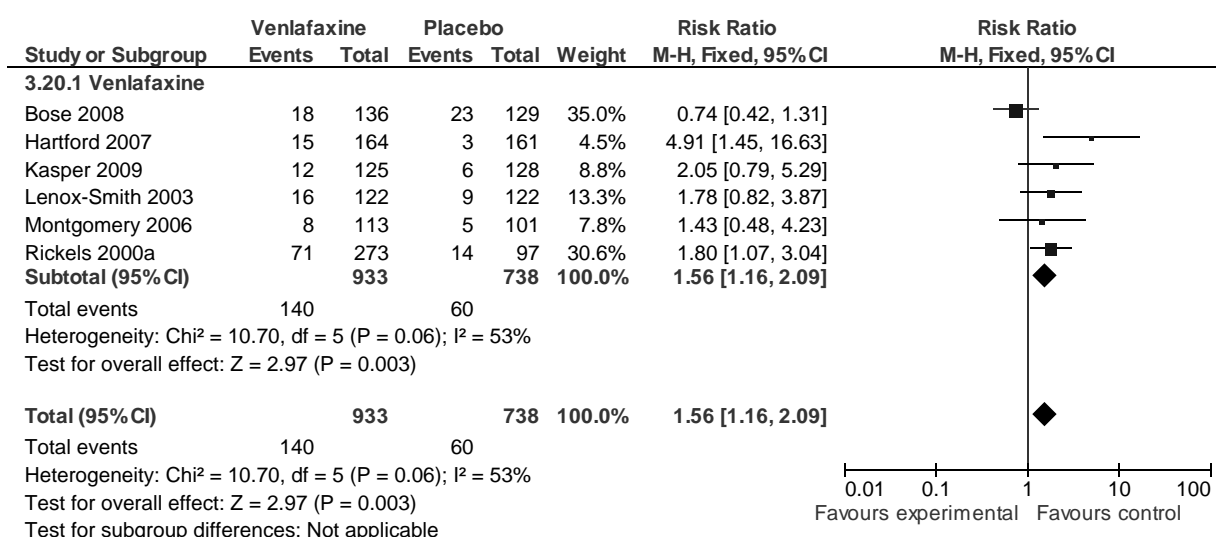
## 3.7 HAM-A



## 3.14 Nausea

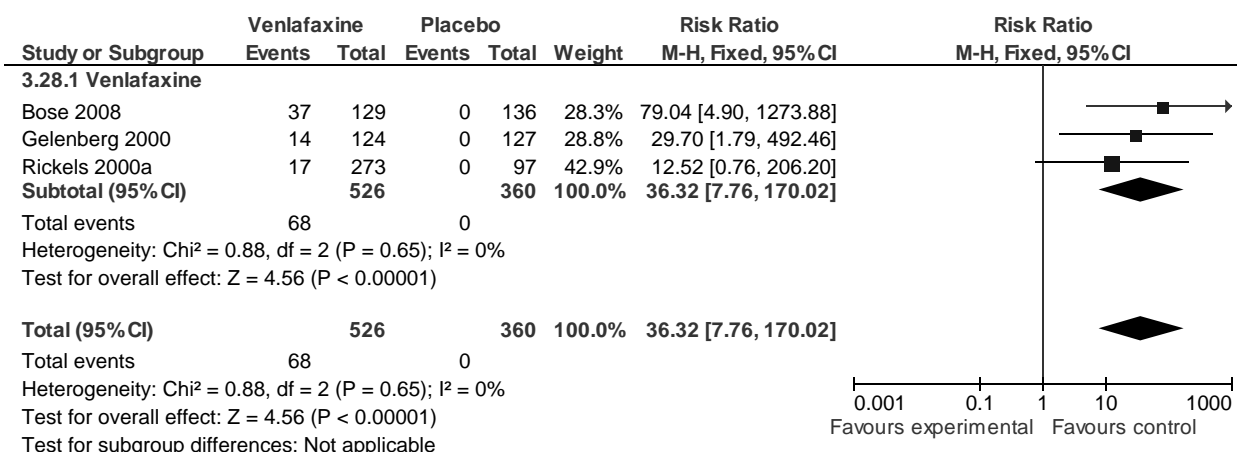


## 3.20 Insomnia



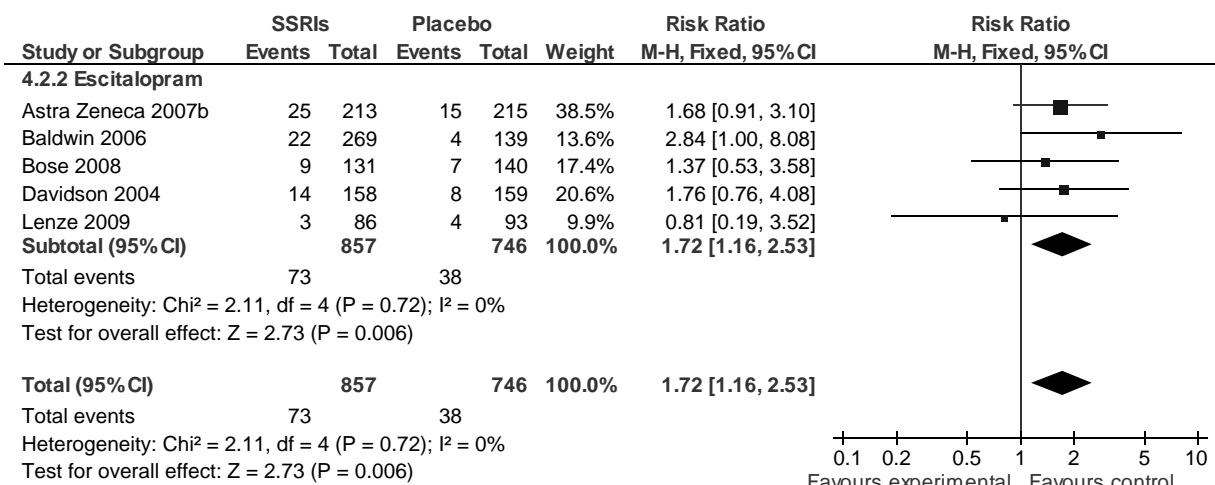
# 1 Antidepressants versus Placebo for GAD

## 3.28 Ejaculation disorder

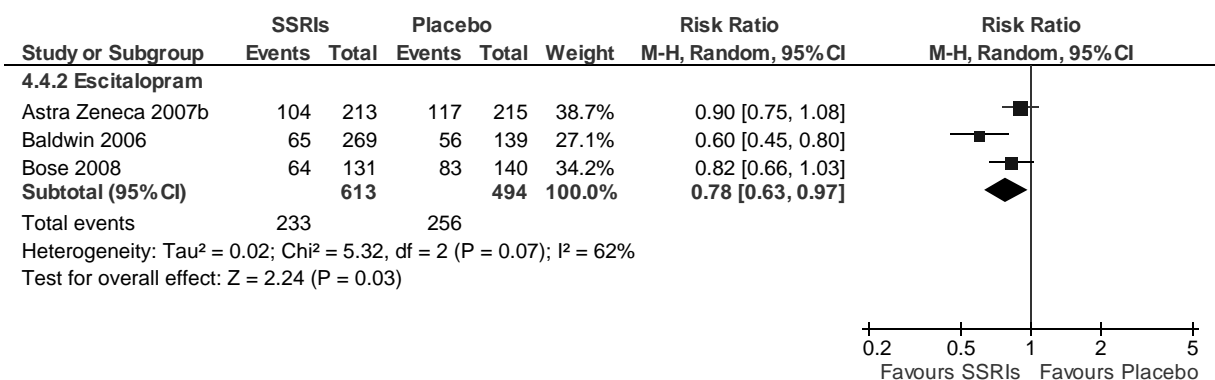


## 4 Escitalopram versus Placebo

### 4.2 Discontinuation due to adverse events

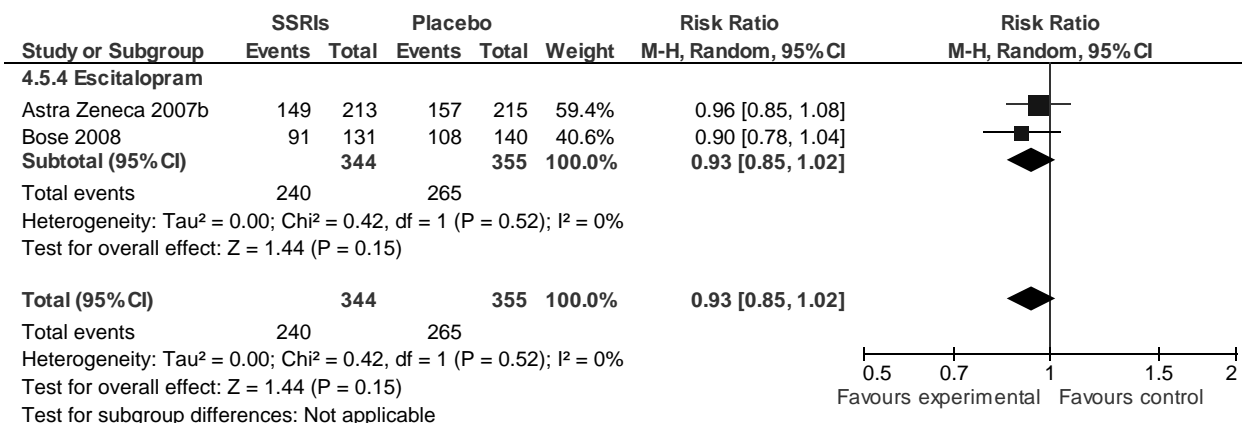


### 4.4 Non-response

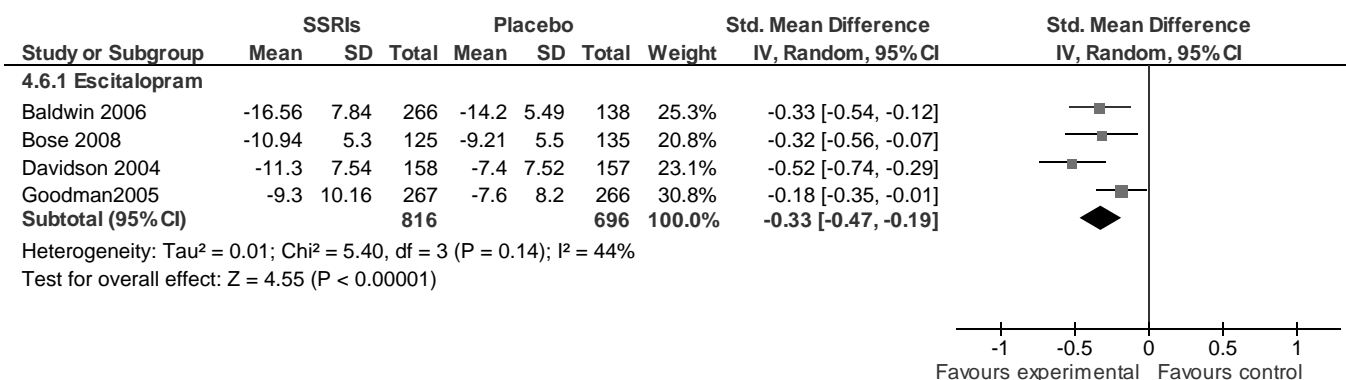


# 1 Antidepressants versus Placebo for GAD

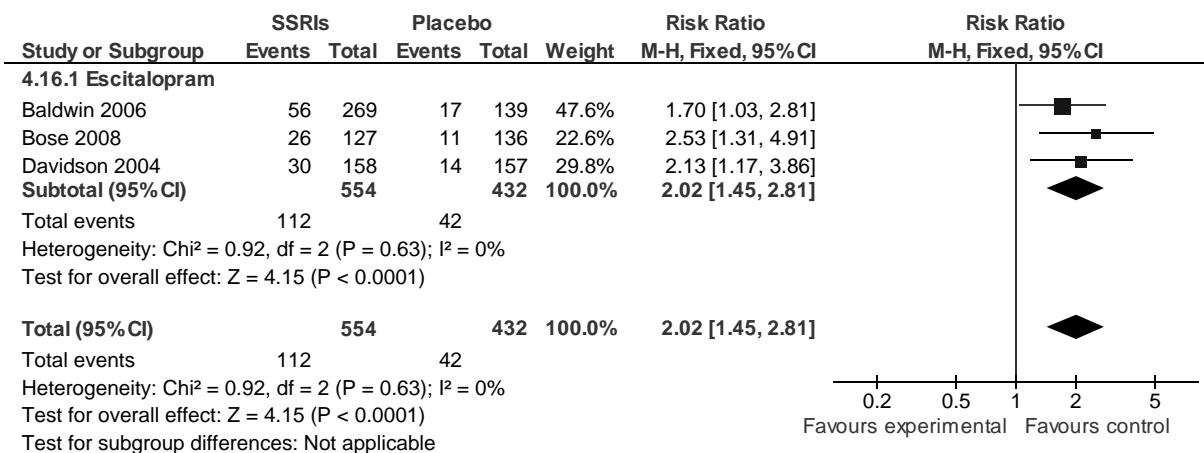
## 4.5 Non-remission



## 4.6 HAM-A (change from baseline)

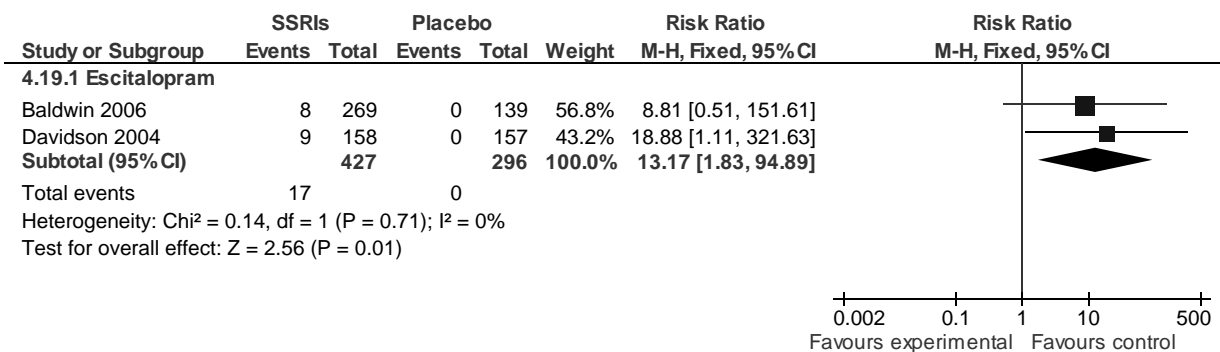


## 4.16 Nausea

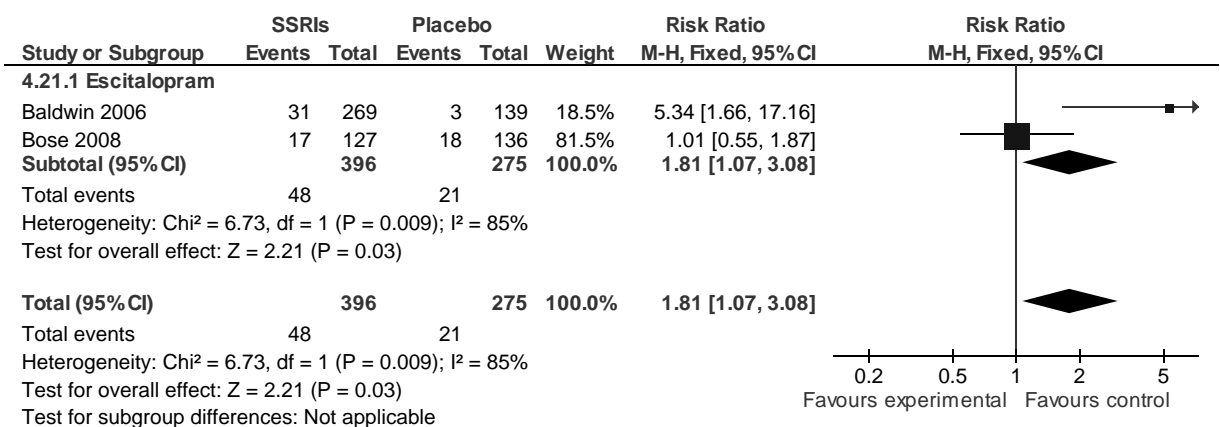


# 1 Antidepressants versus Placebo for GAD

## 4.19 Anorgasmia

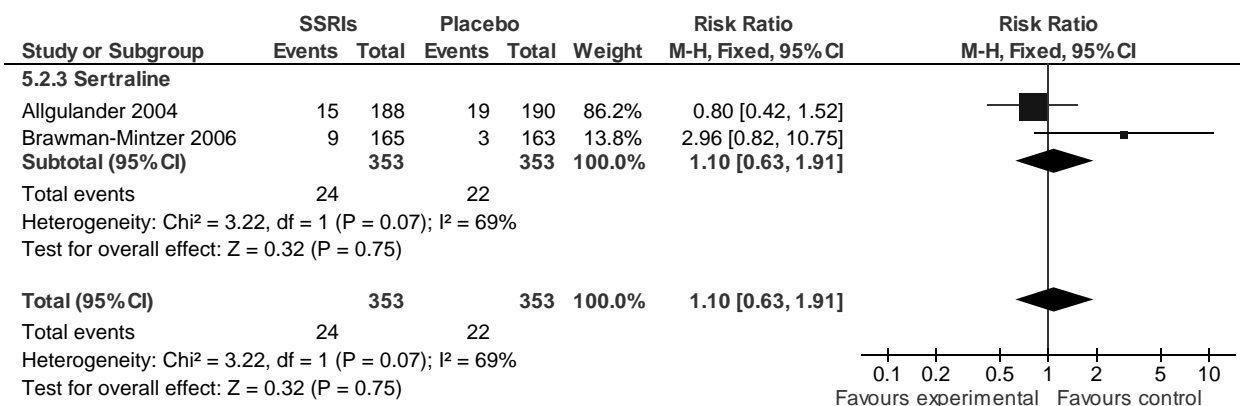


## 4.21 Insomnia



## 5 Sertraline versus Placebo

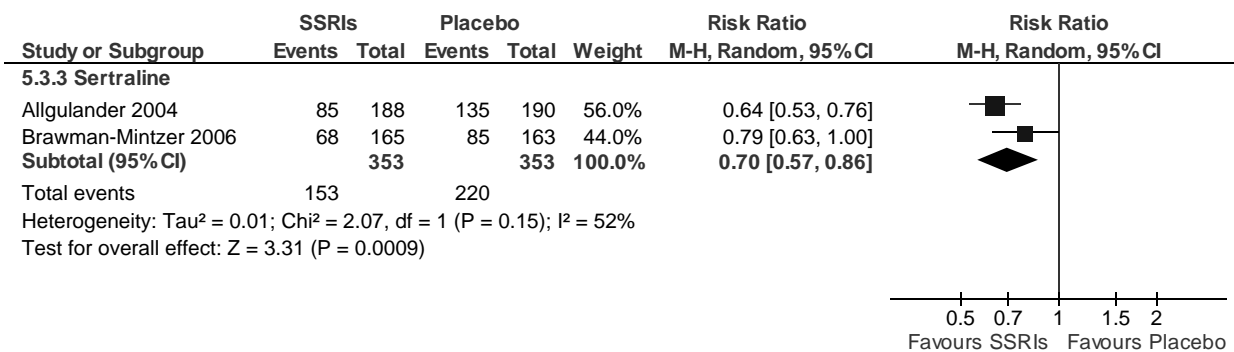
### 5.2 Discontinuation due to adverse events



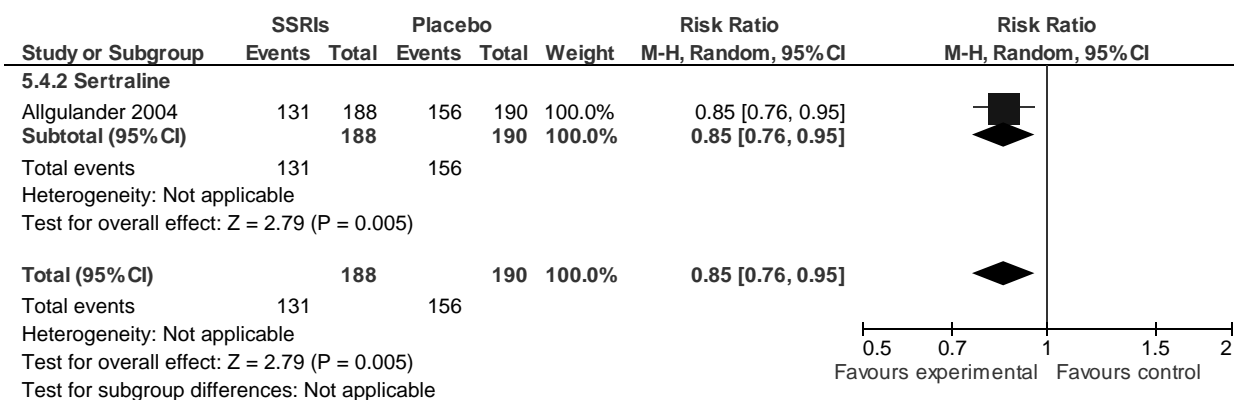


# 1 Antidepressants versus Placebo for GAD

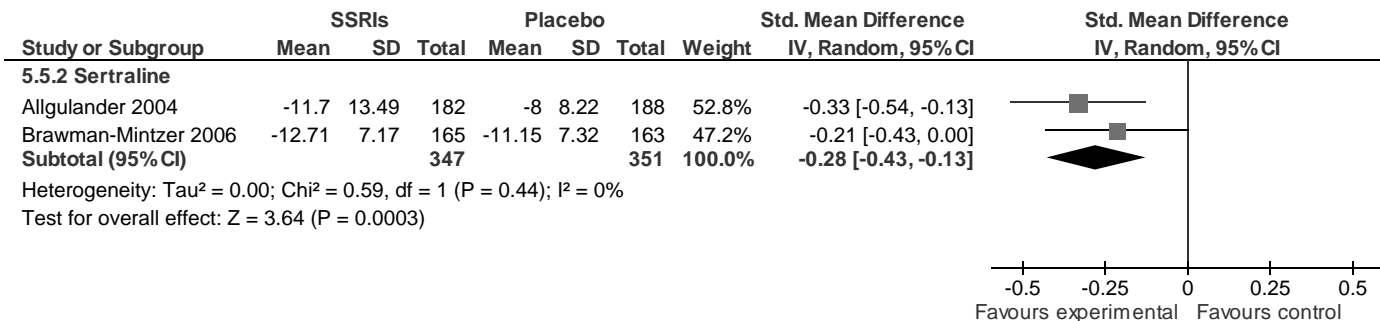
## 5.3 Non-response



## 5.4 Non-remission

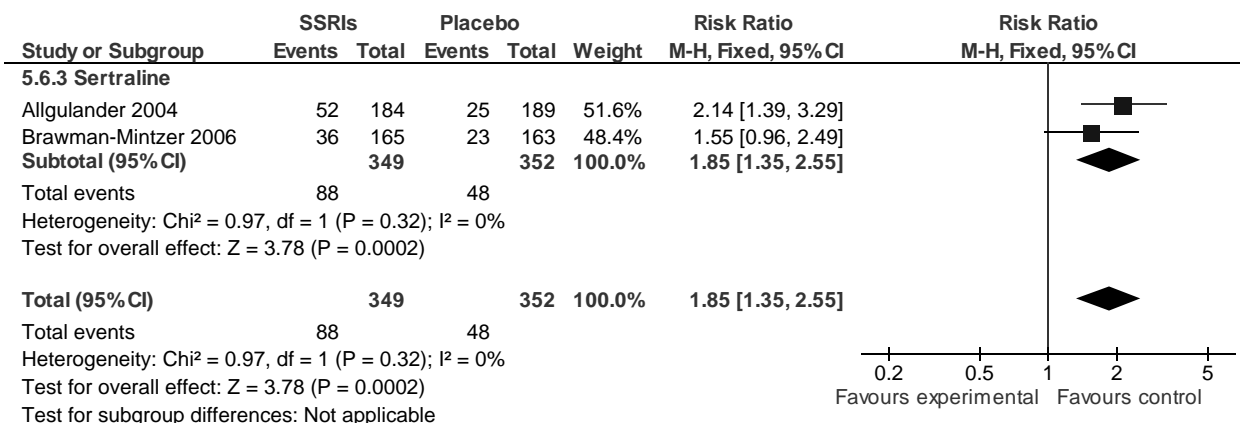


## 5.5 HAM-A (change from baseline)

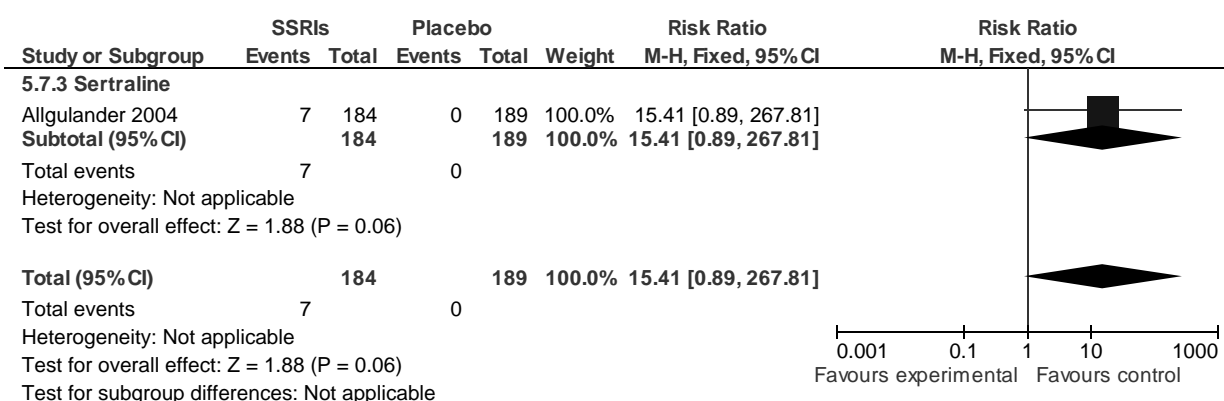


# 1 Antidepressants versus Placebo for GAD

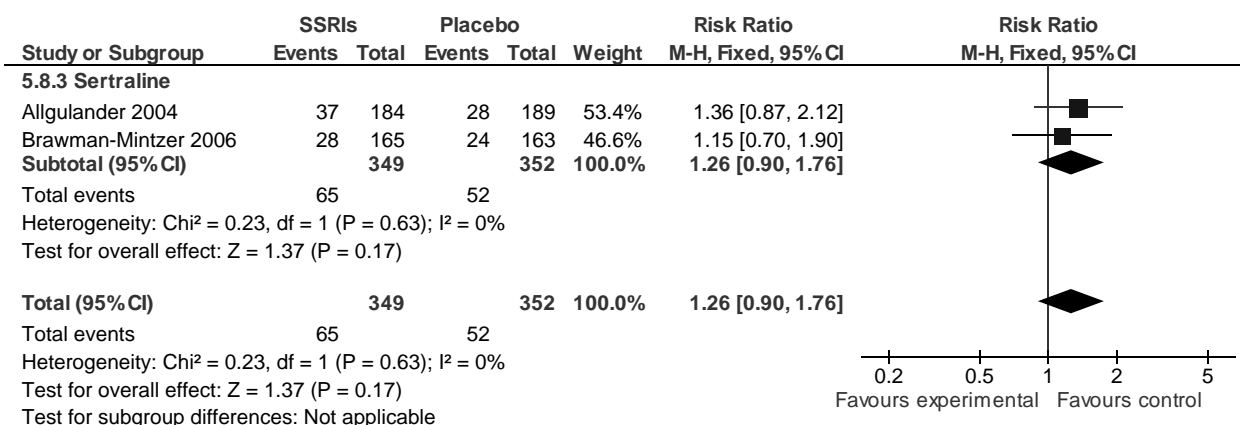
## 5.6 Nausea



## 5.7 Ejaculation disorder



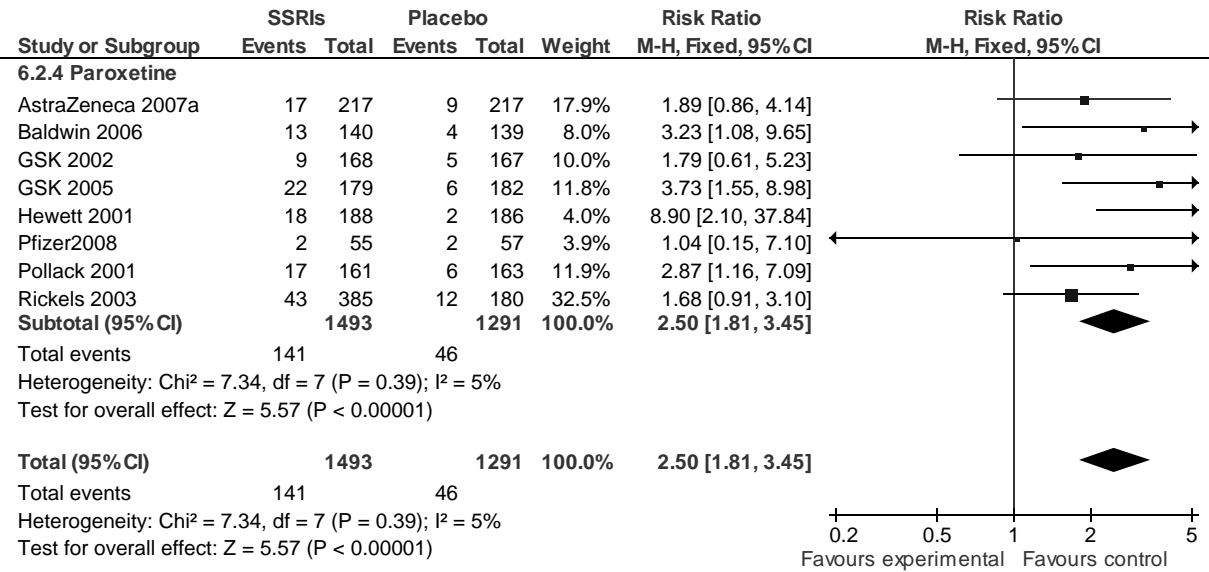
## 5.8 Insomnia



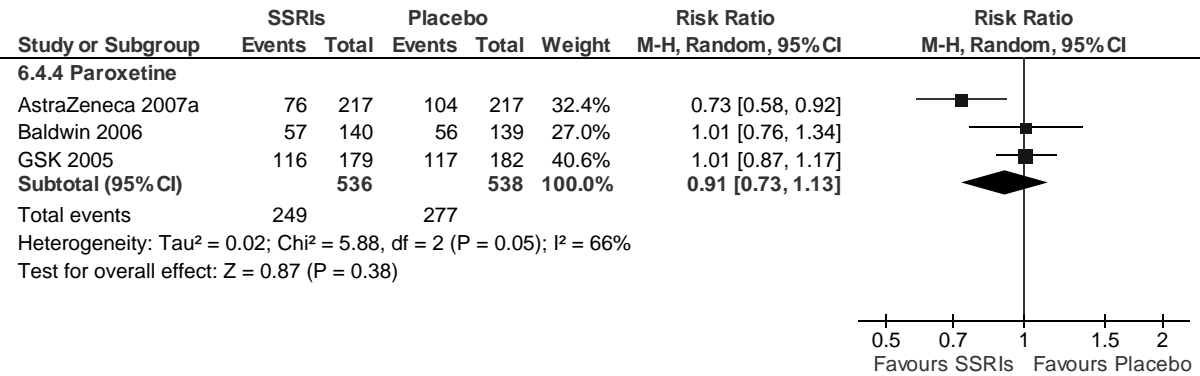
## 6 Paroxetine versus Placebo

# 1 Antidepressants versus Placebo for GAD

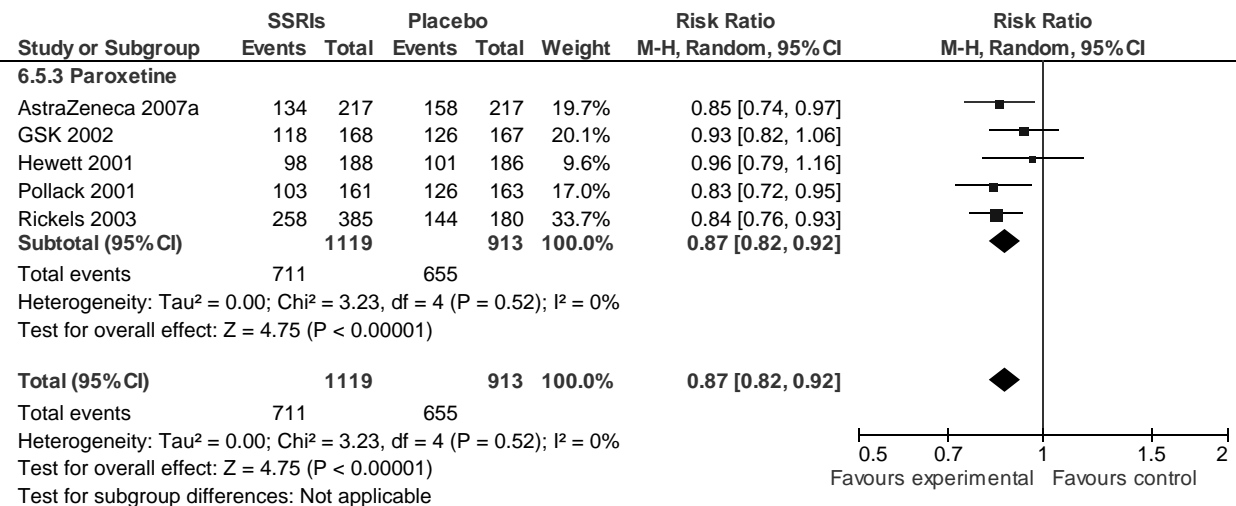
## 6.2 Discontinuation due to adverse events



## 6.4 Non-response

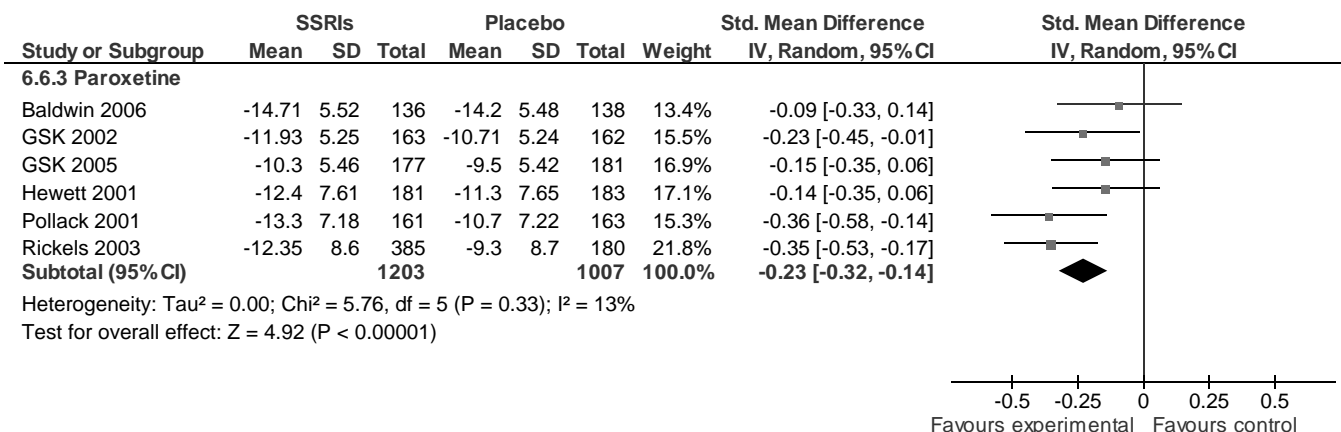


## 6.5 Non-remission

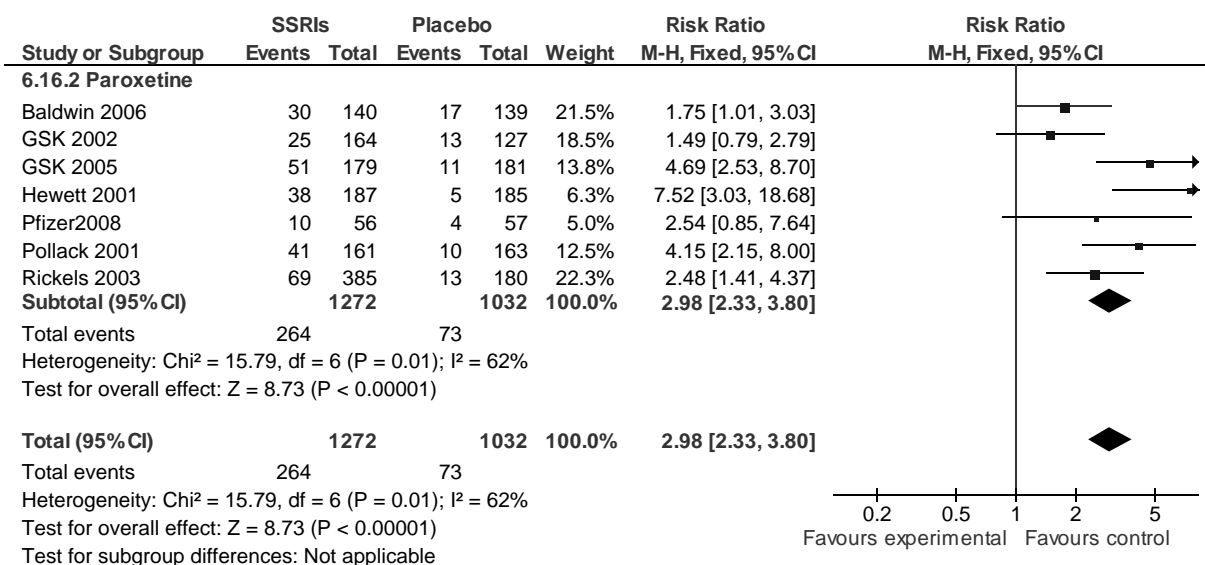


# 1 Antidepressants versus Placebo for GAD

## 6.6 HAM-A (change from baseline)

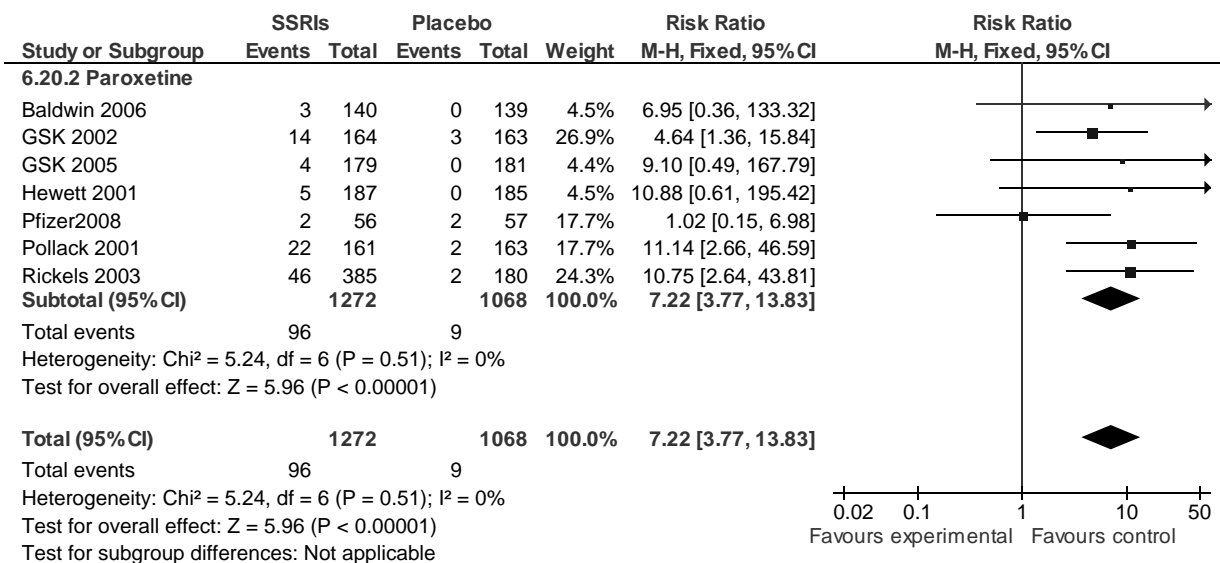


## 6.16 Nausea

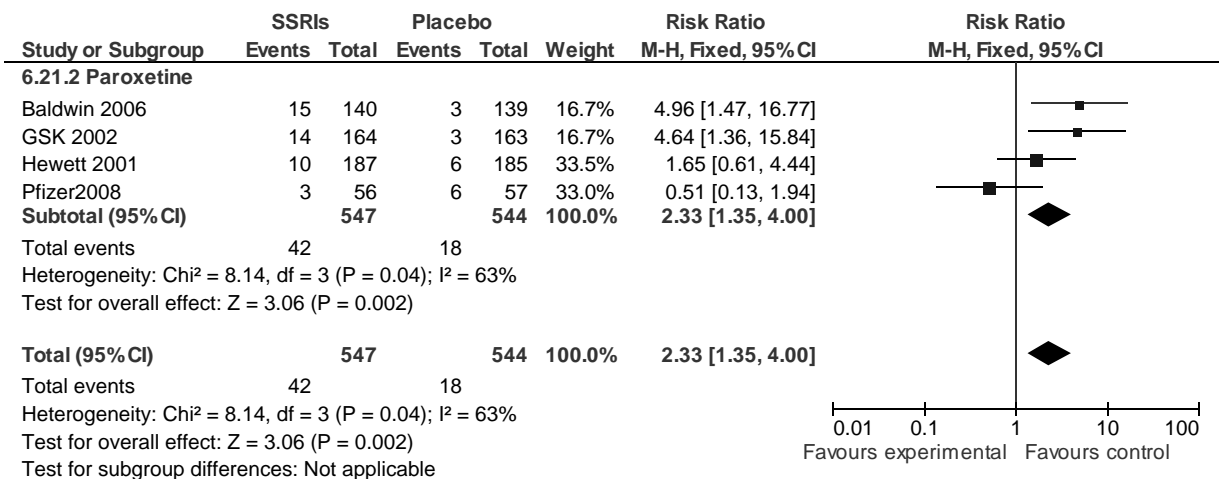


# 1 Antidepressants versus Placebo for GAD

## 6.20 Sexual problem



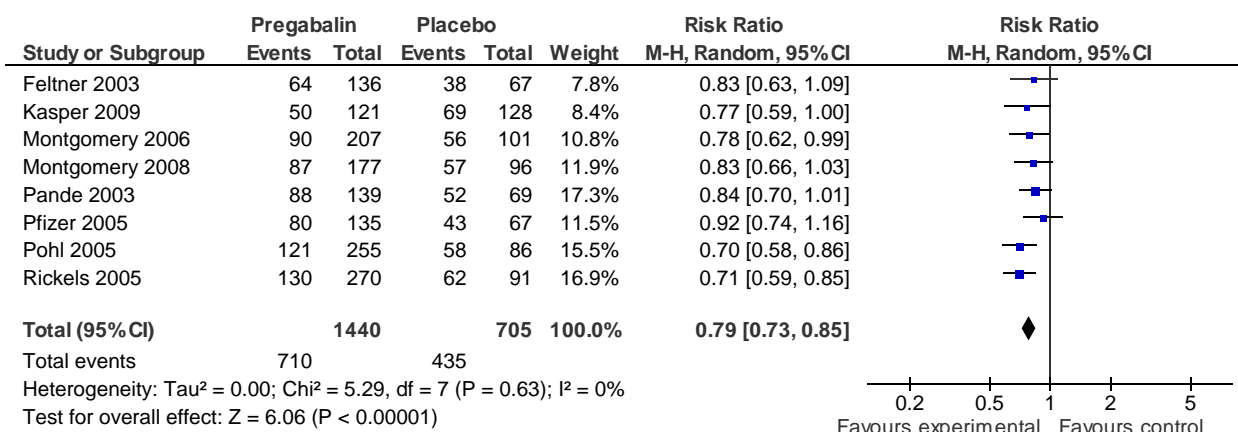
## 6.21 Insomnia



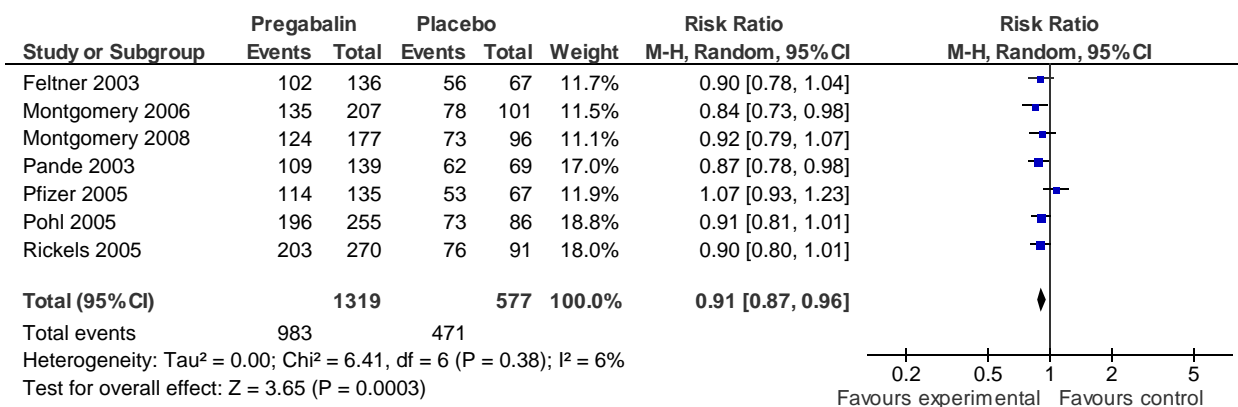
## 7 Pregabalin versus Placebo

## 7 Pregabalin vs Placebo

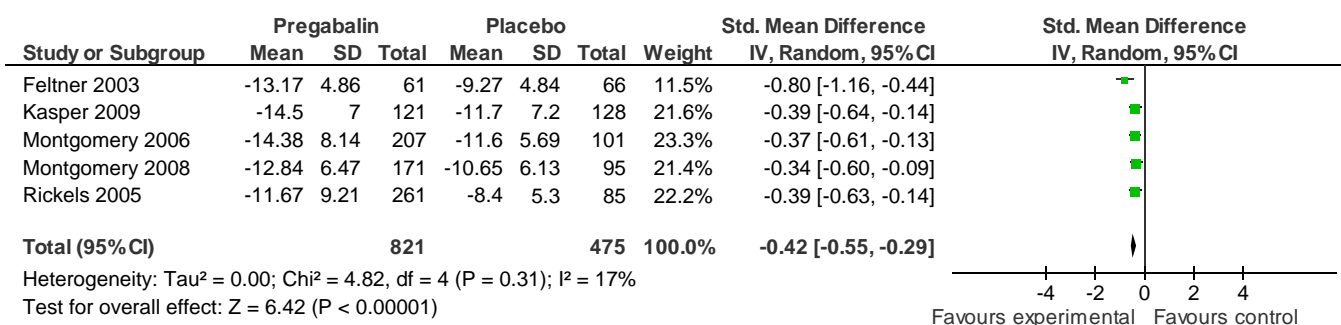
### 7.3 Non-response



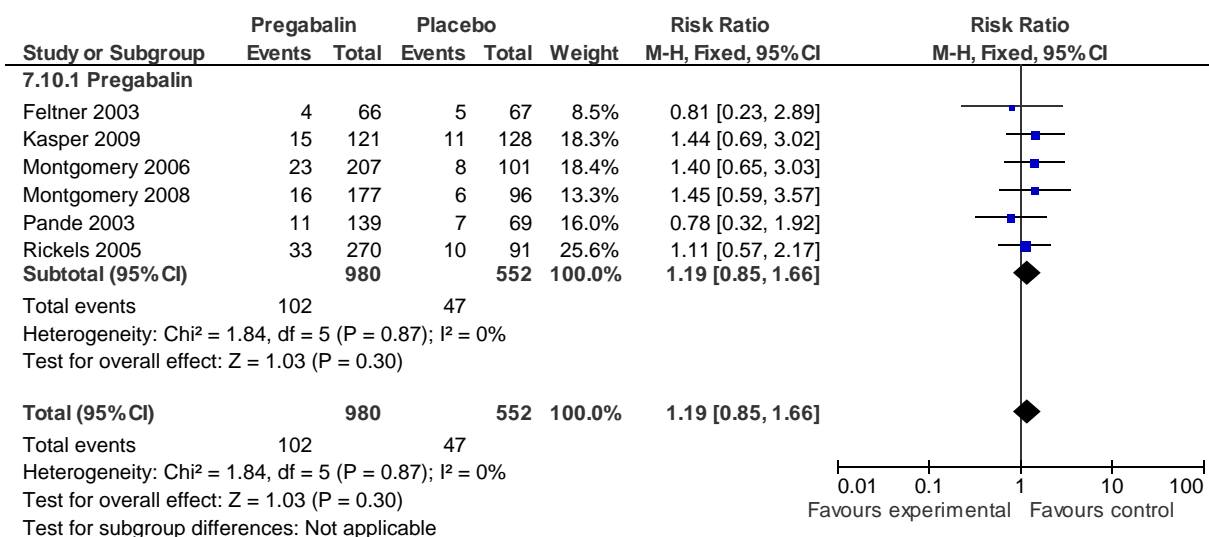
### 7.4 Non-remission



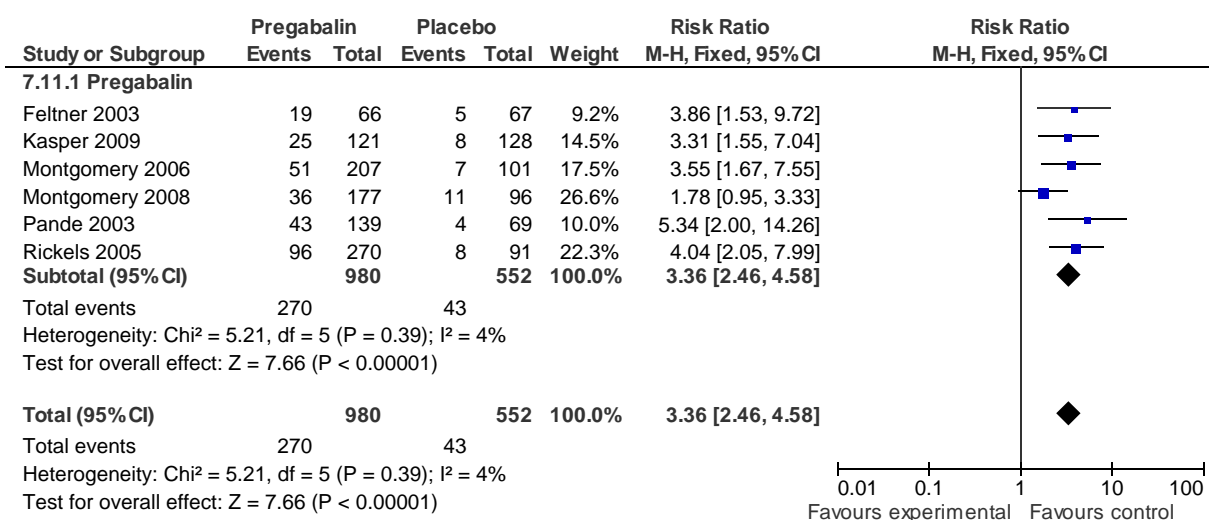
### 7.5 HAM-A



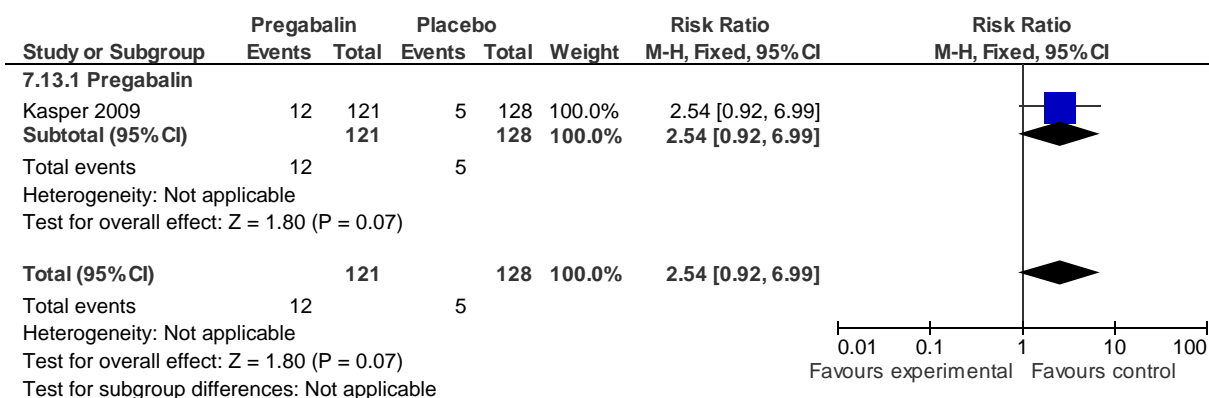
### 7.10 Nausea



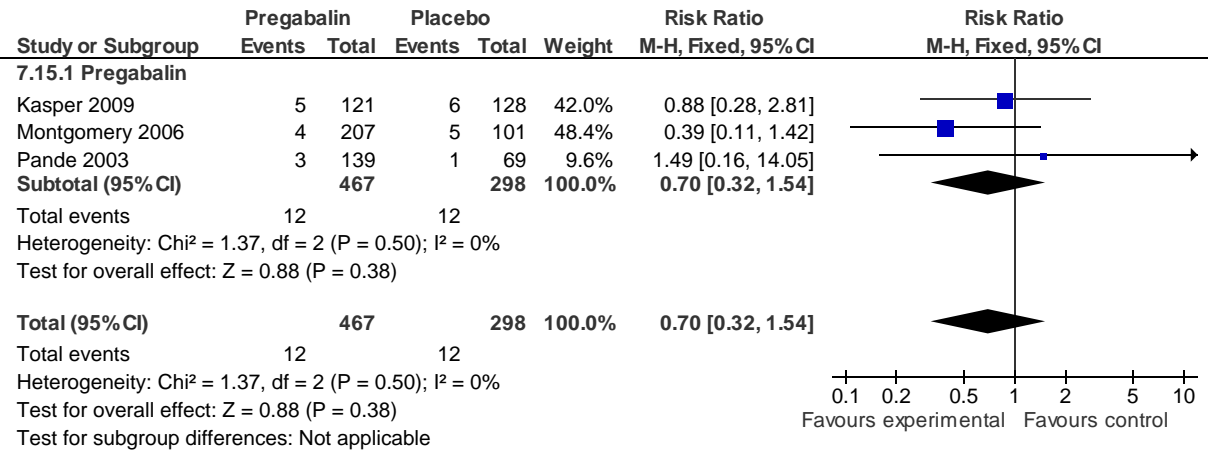
### 7.11 Dizziness



### 7.13 Fatigue



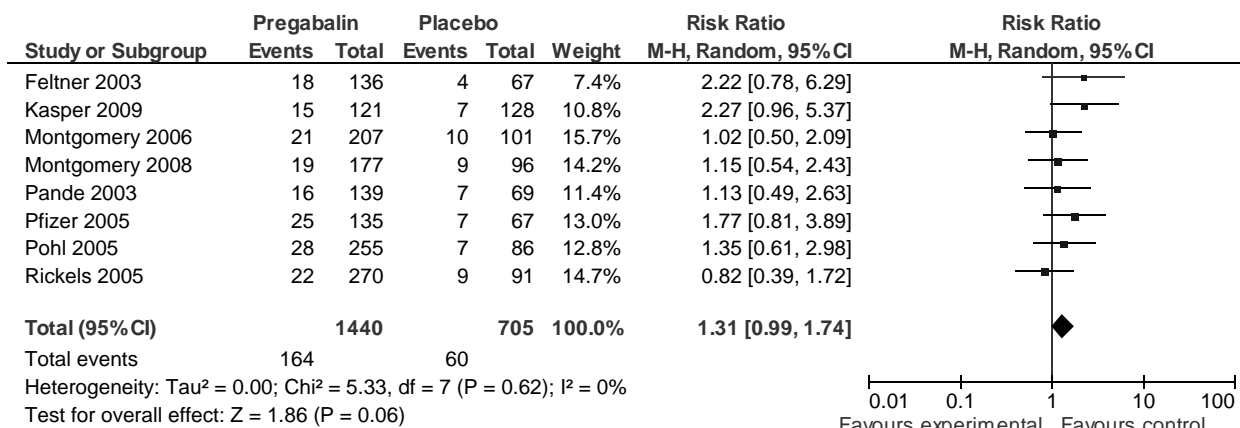
7.15 Insomnia





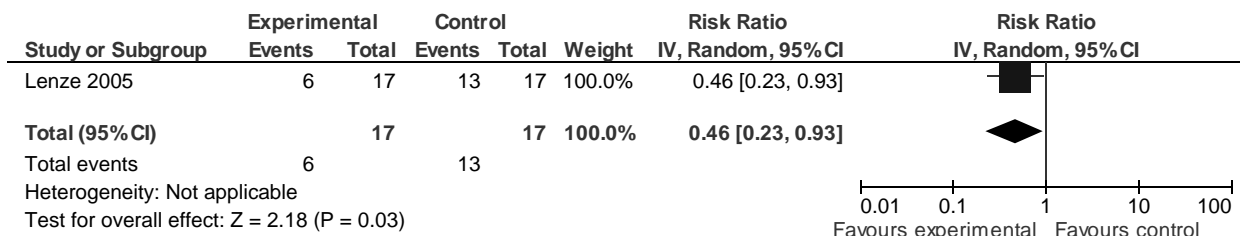
# 1 Antidepressants versus Placebo for GAD

## 7.2 Discontinuation due to adverse events

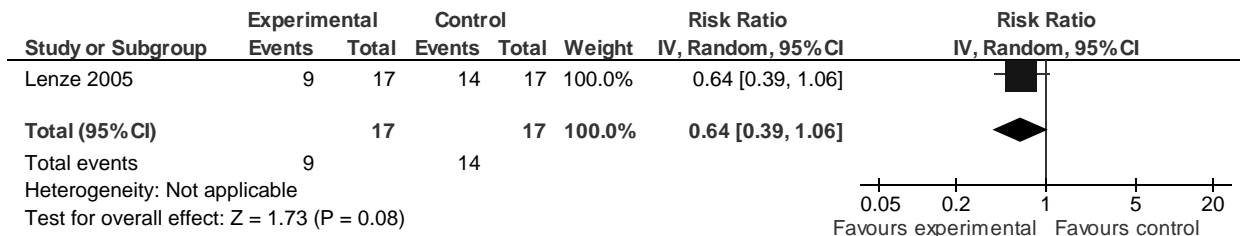


## 35 Citalopram versus Placebo

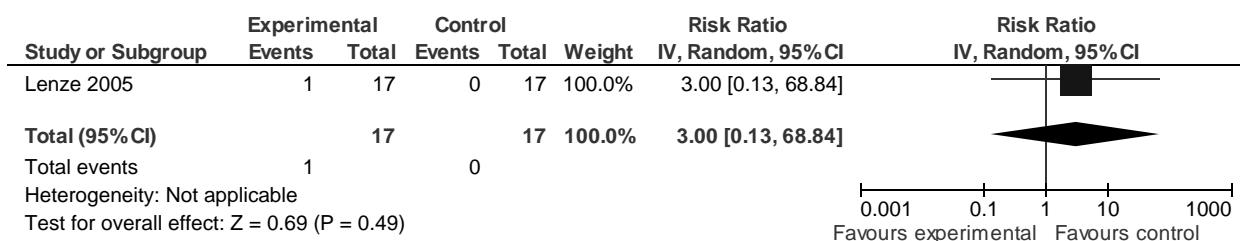
### 35.1 Non-response



### 35.2 Non-remission

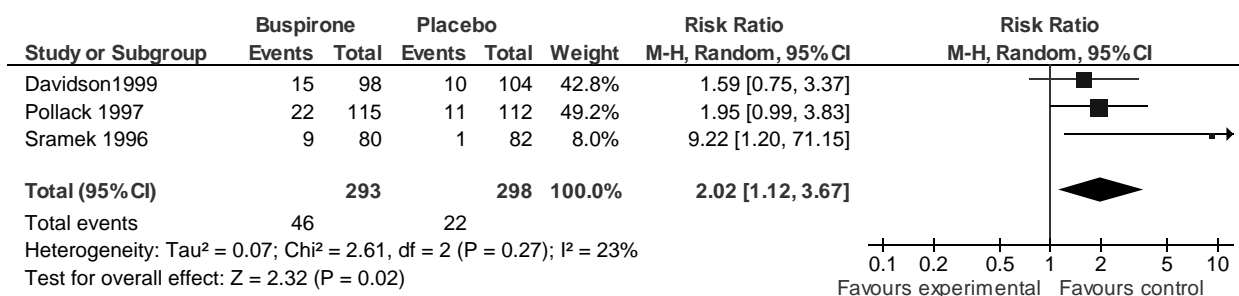


### 35.3 Discontinuation due to adverse events

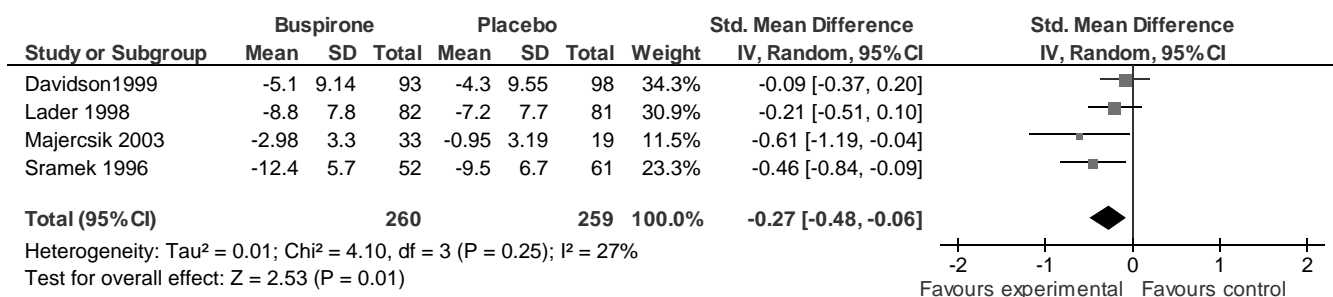


## 8 Buspirone vs Placebo

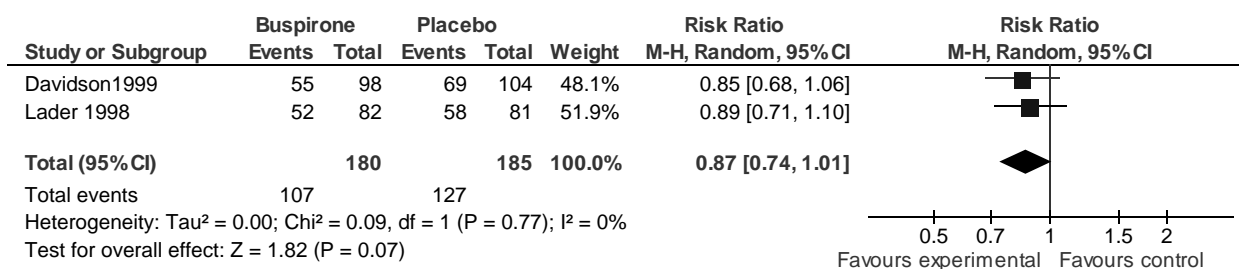
### 8.2 Discontinuation due to adverse events



### 8.3 HAM-A

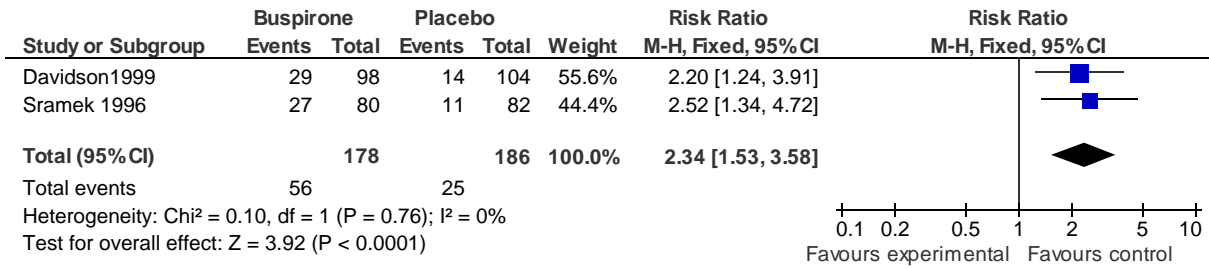


### 8.4 Non-response

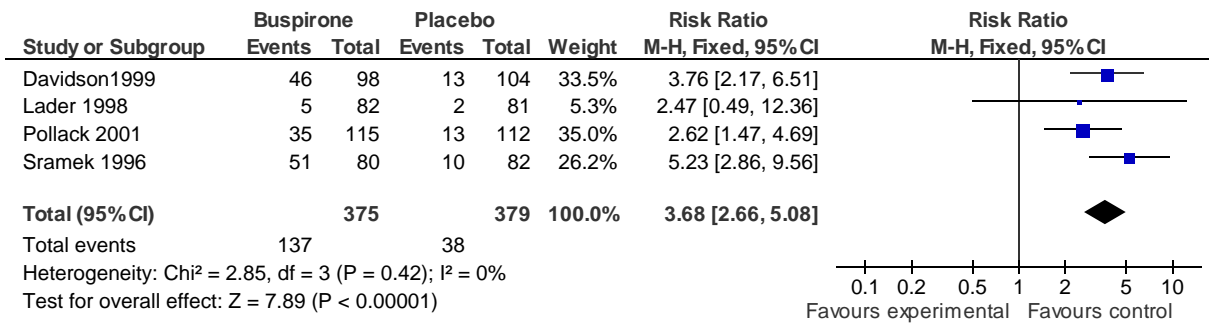


## 2 Benzodiazepine versus Placebo for GAD

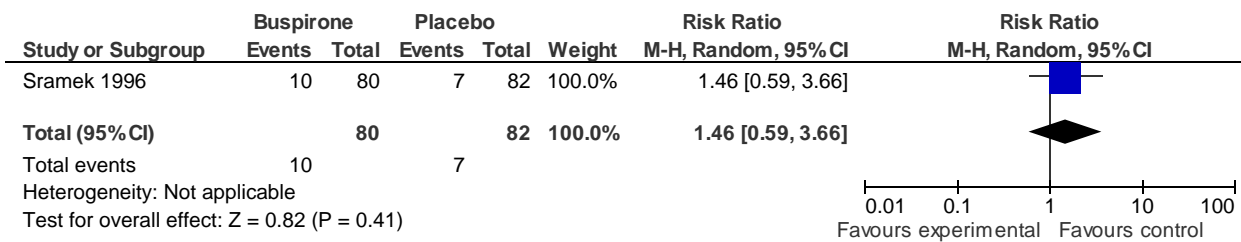
### 8.5 Nausea



### 8.6 Dizziness



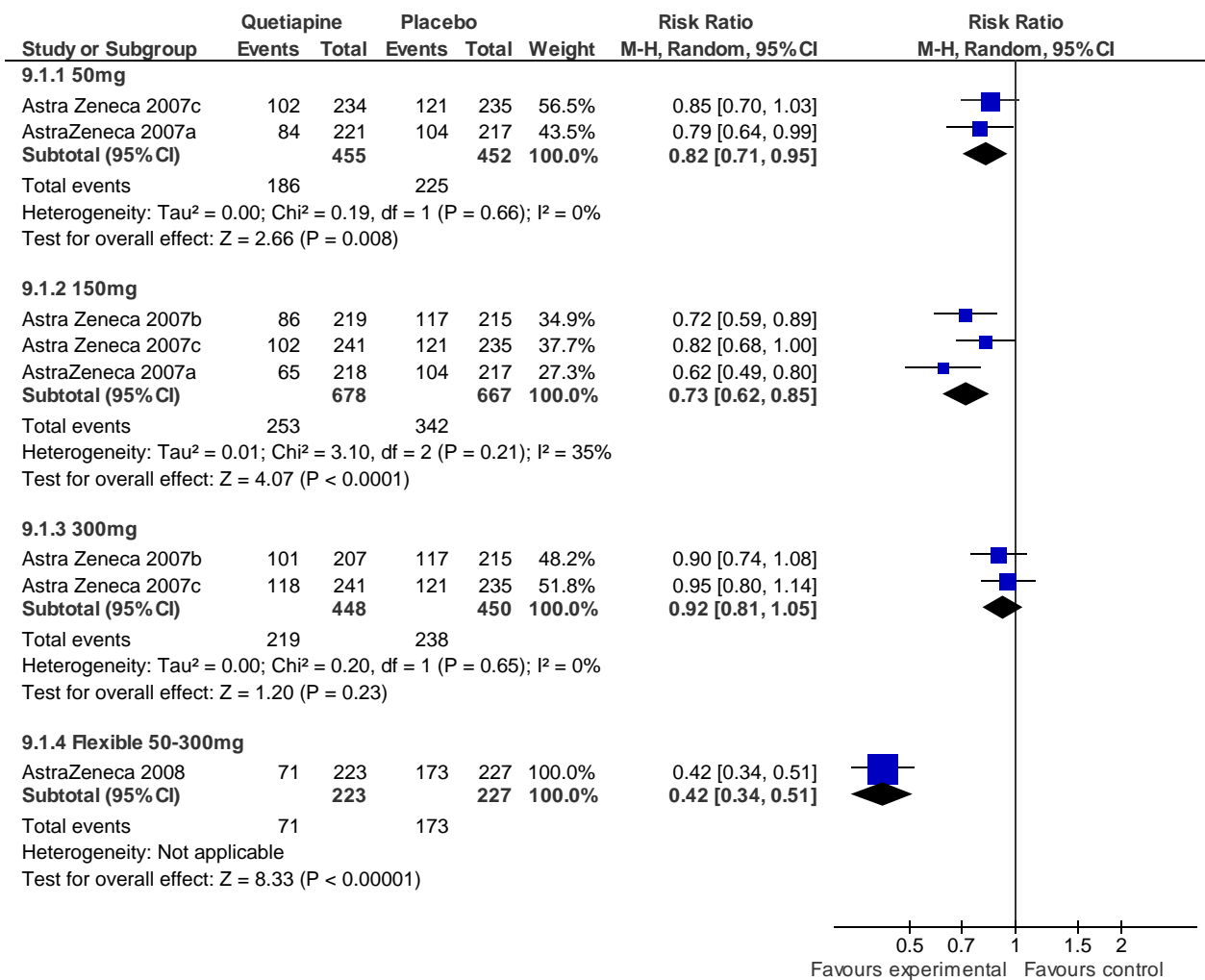
### 8.11 Insomnia



### 9 Quetiapine versus Placebo

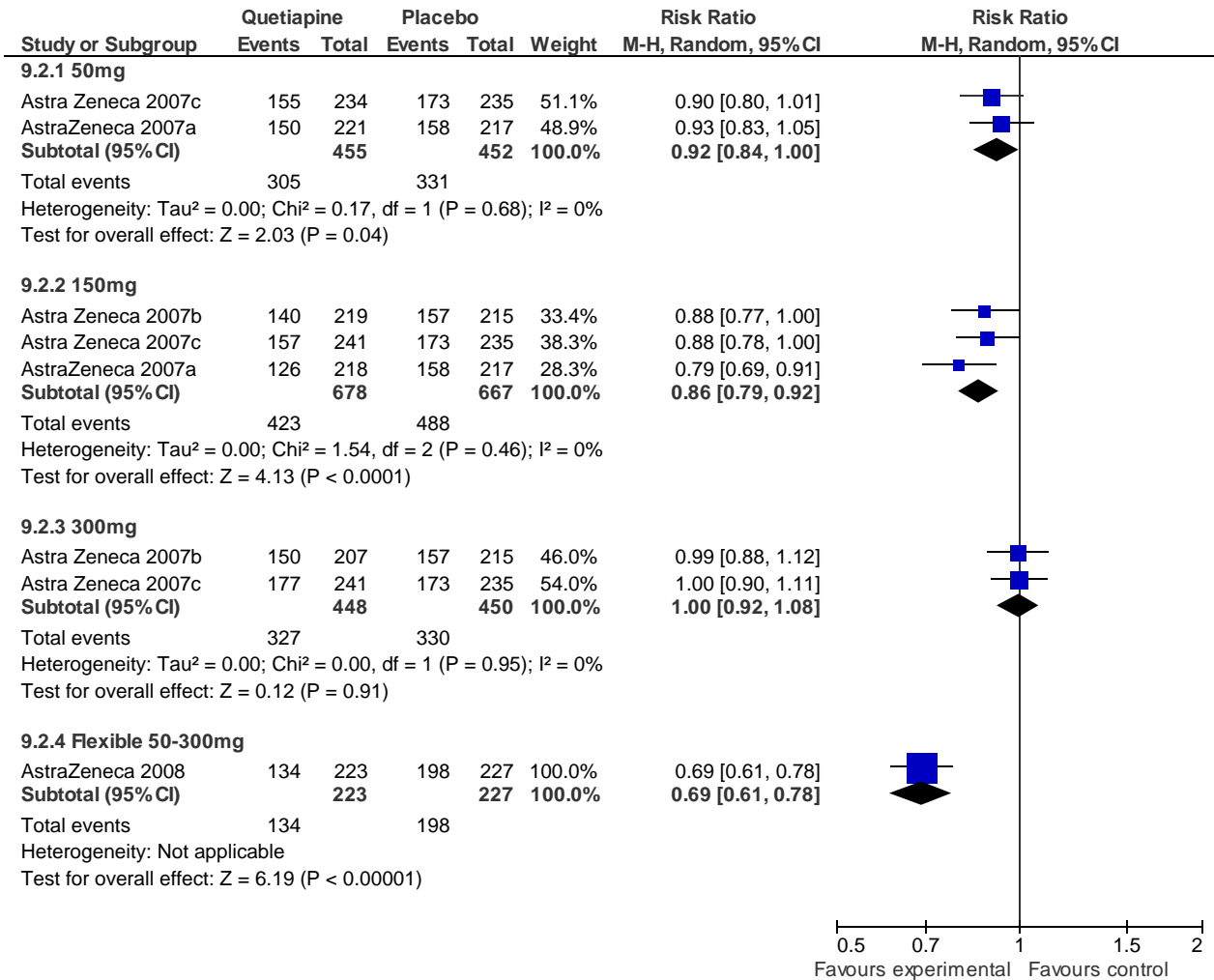
## 2 Benzodiazepine versus Placebo for GAD

### 9.1 Non-response



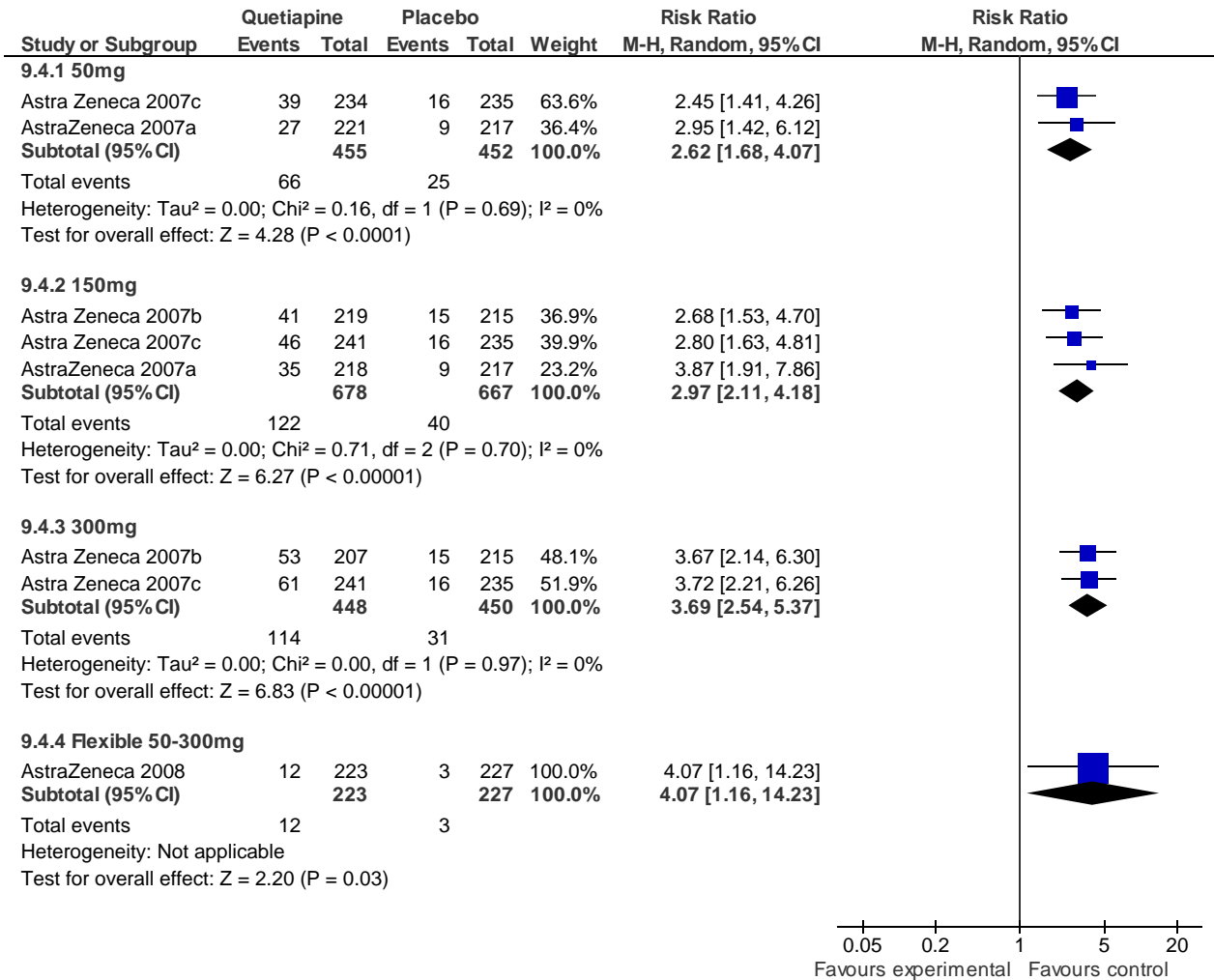
## 2 Benzodiazepine versus Placebo for GAD

### 9.2 Non-remission



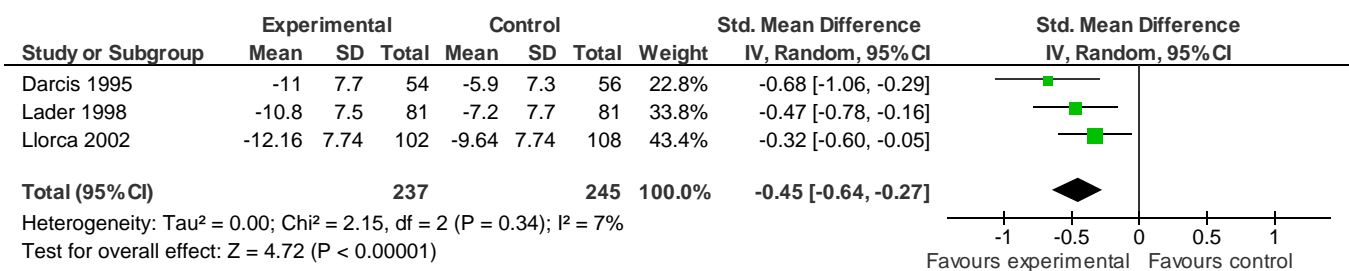
## 2 Benzodiazepine versus Placebo for GAD

### 9.4 Discontinuation due to adverse events



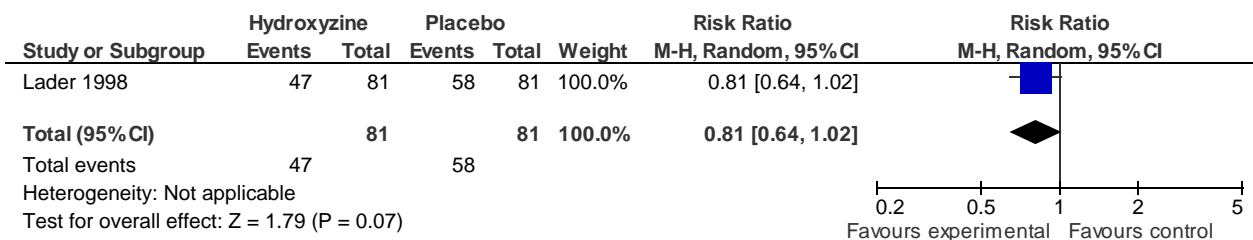
### 10 Hydroxyzine versus Placebo

#### 10.1 HAM-A

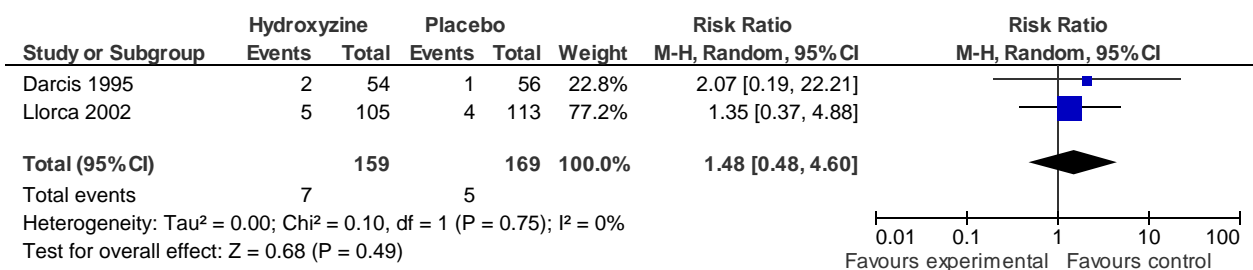


## 2 Benzodiazepine versus Placebo for GAD

### 10.2 Non-response

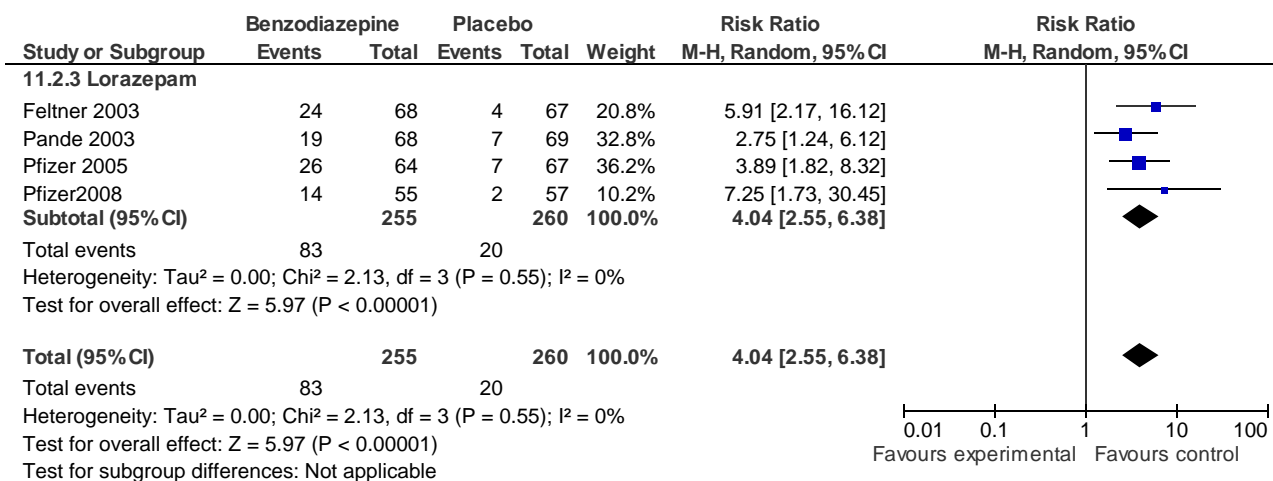


### 10.4 Discontinuation due to adverse events



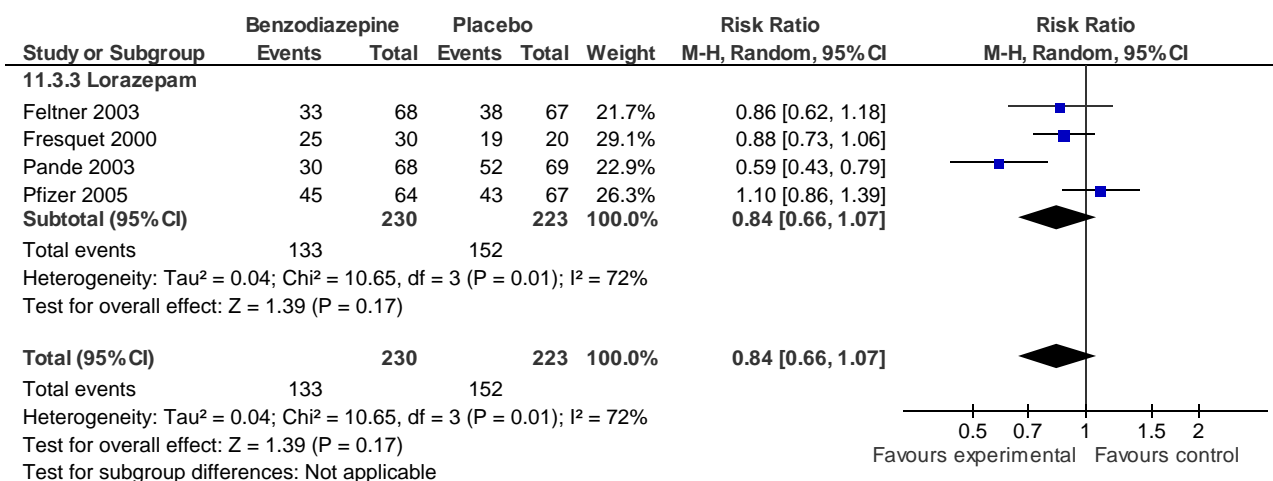
## 11 Lorazepam versus Placebo

### 11.2 Discontinuation due to adverse events

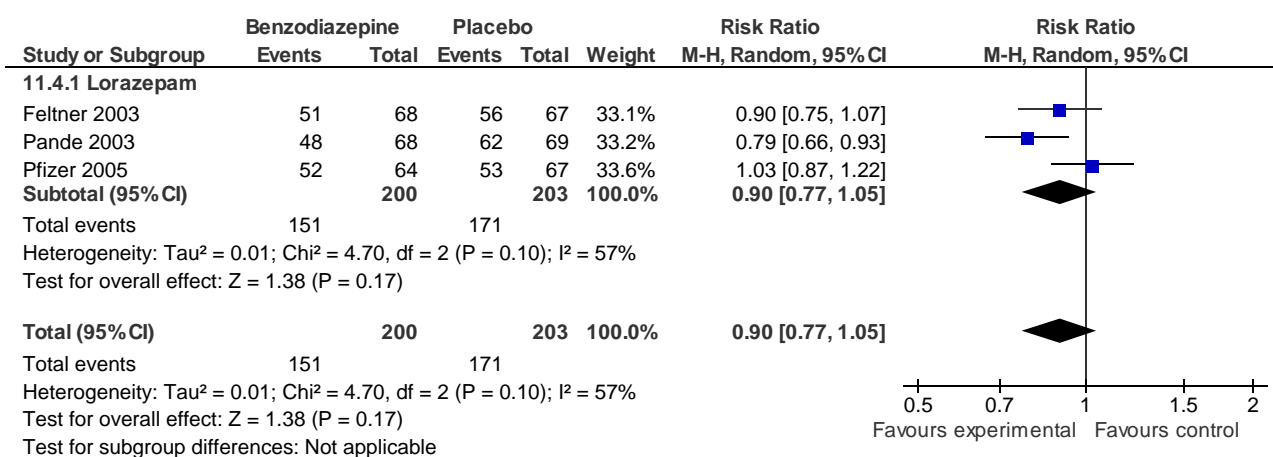


## 2 Benzodiazepine versus Placebo for GAD

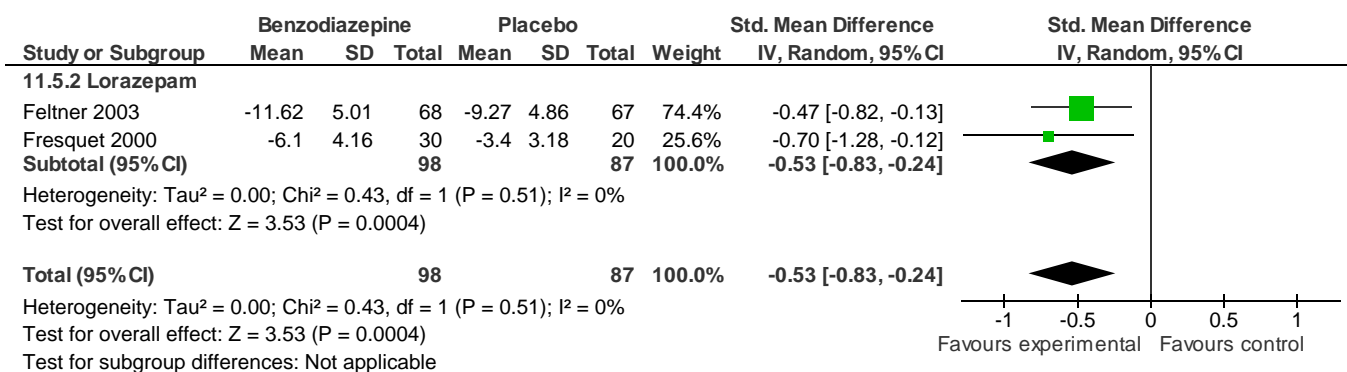
### 11.3 Non-response



### 11.4 Non-remission



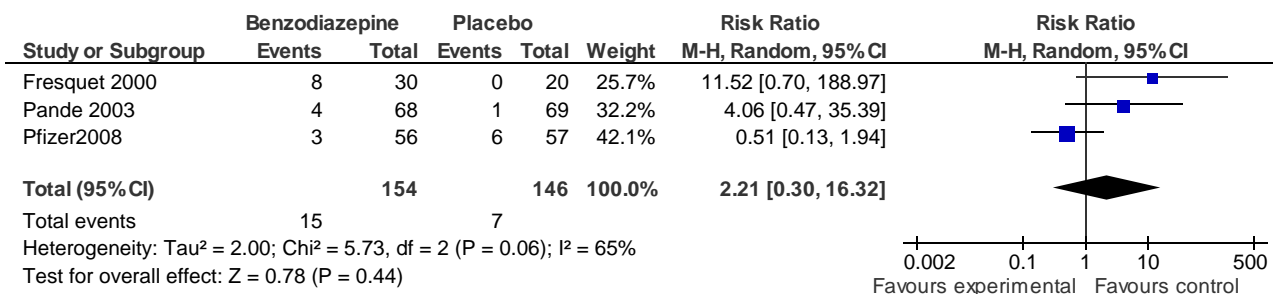
### 11.5 HAM-A



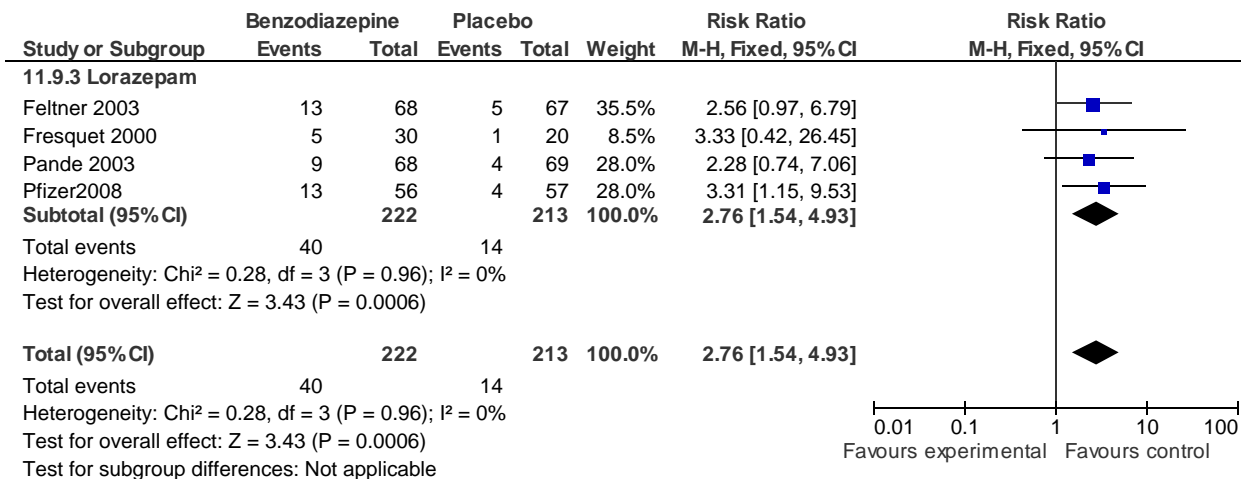


## 2 Benzodiazepine versus Placebo for GAD

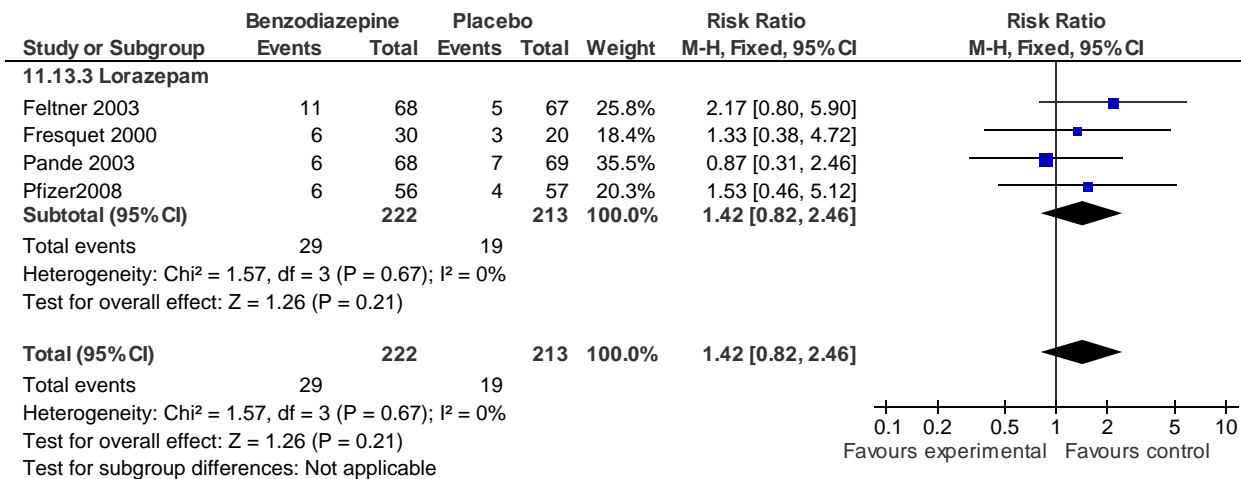
### 11.7 Insomnia



### 11.9 Dizziness



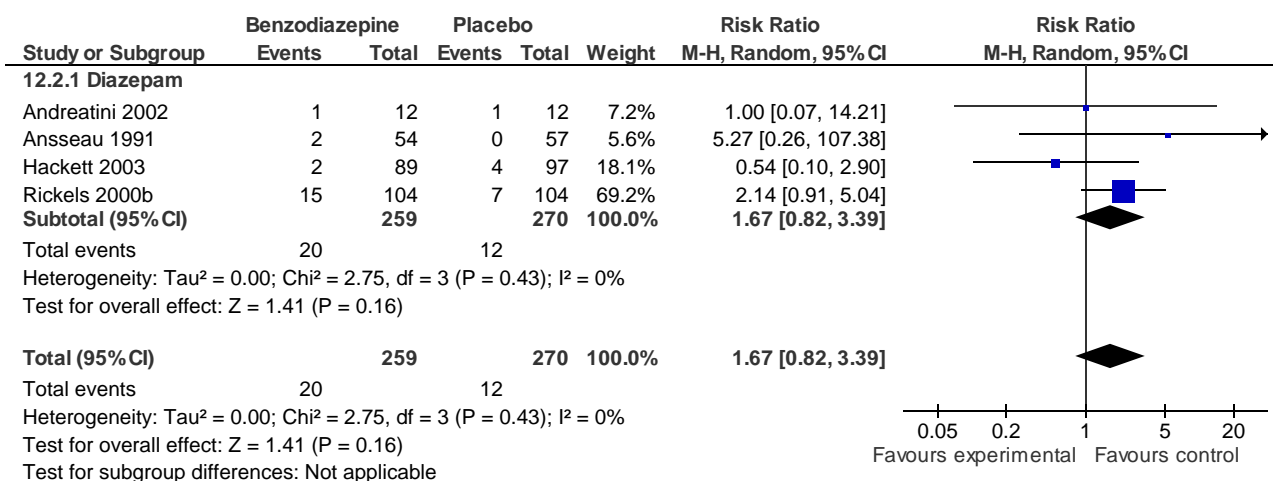
### 11.13 Nausea



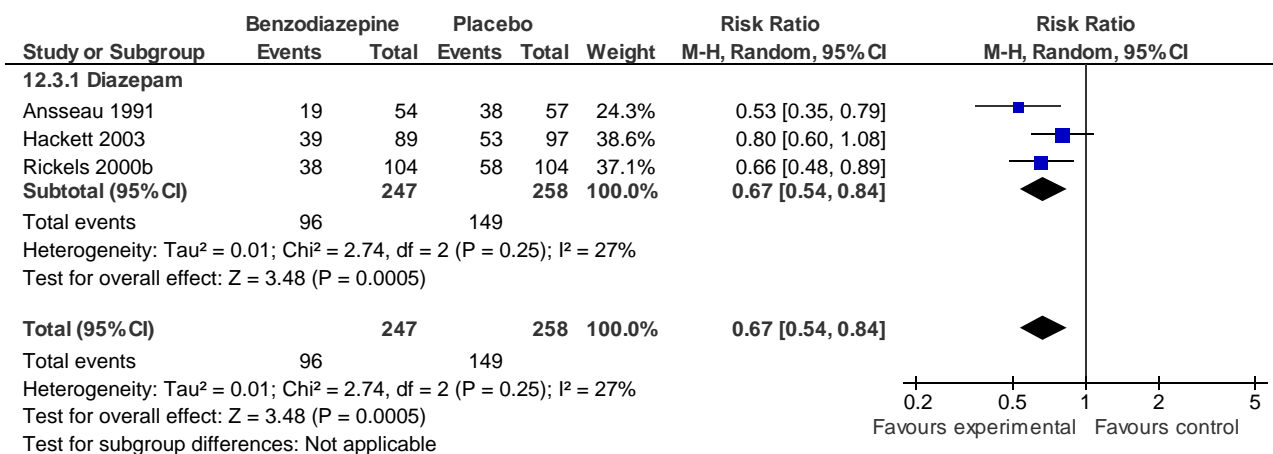
## 12 Diazepam versus Placebo

## 2 Benzodiazepine versus Placebo for GAD

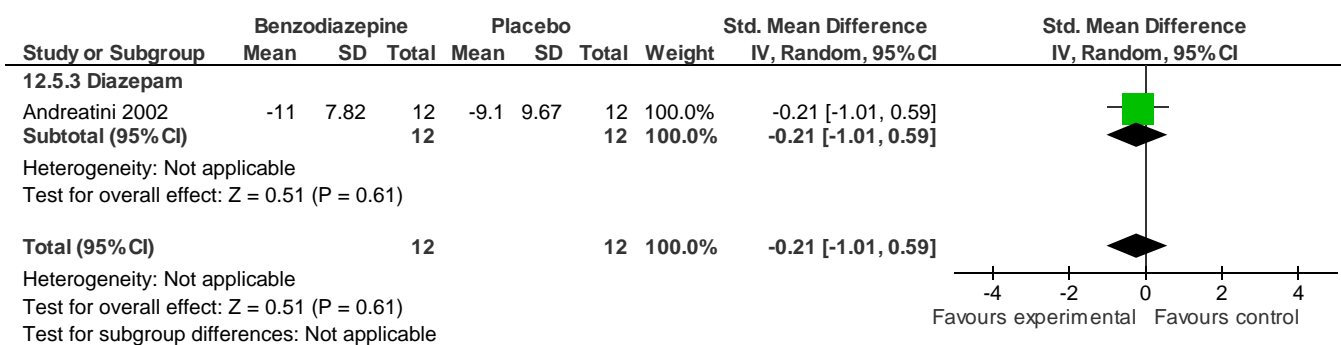
### 12.2 Discontinuation due to adverse events



### 12.3 Non-response

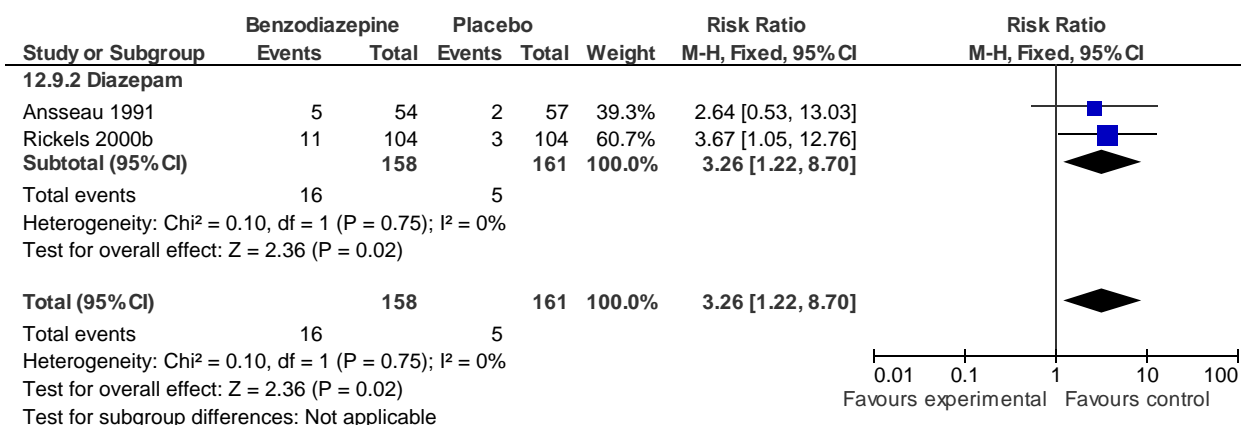


### 12.5 HAM-A

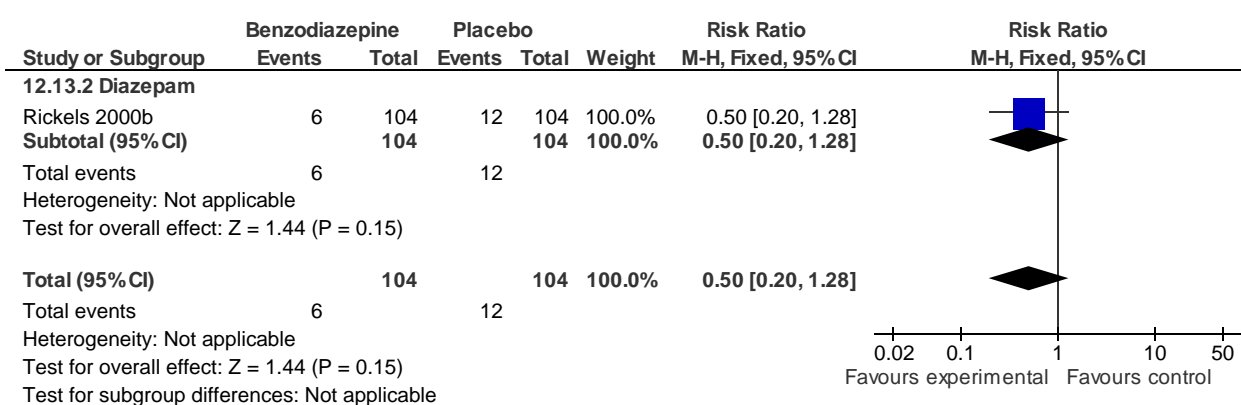


## 2 Benzodiazepine versus Placebo for GAD

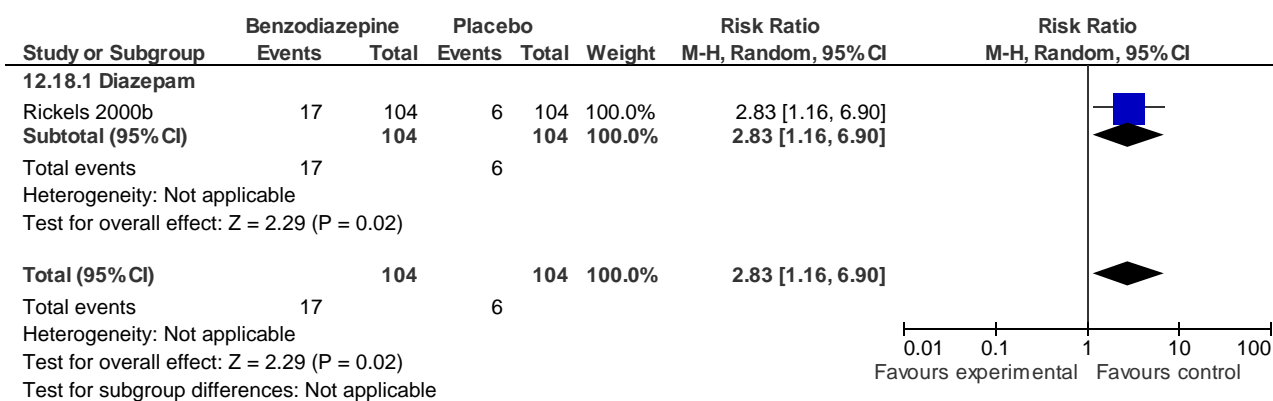
### 12.9 Dizziness



### 12.13 Nausea

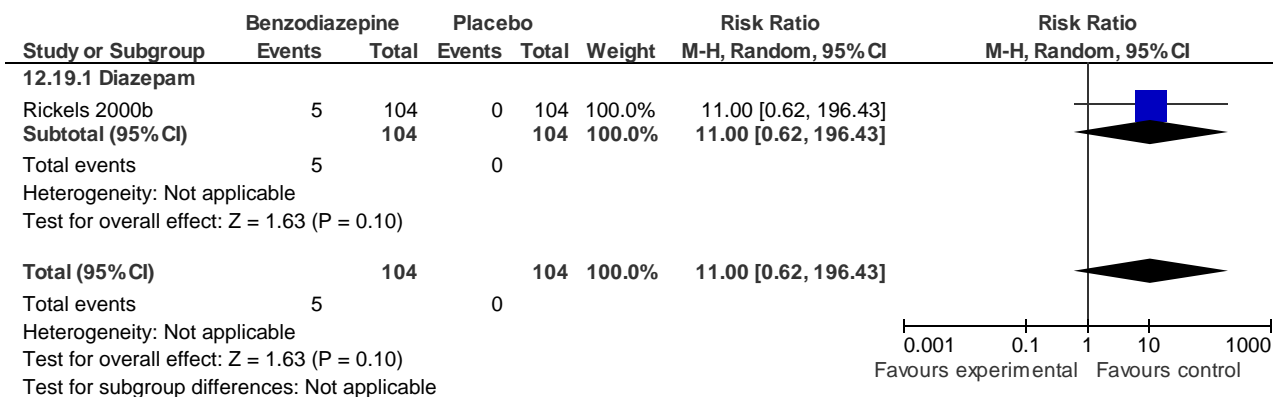


### 12.18 Fatigue



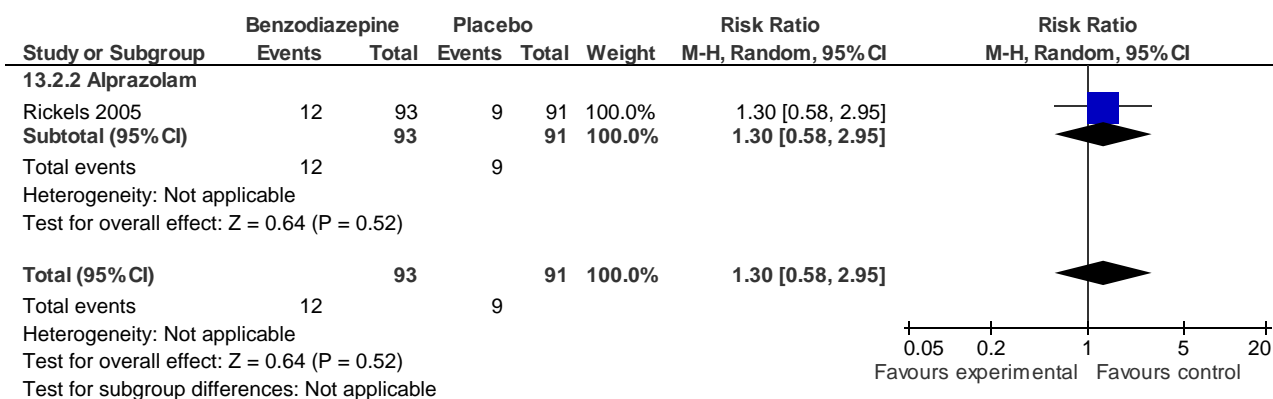
## 2 Benzodiazepine versus Placebo for GAD

### 12.19 Libido

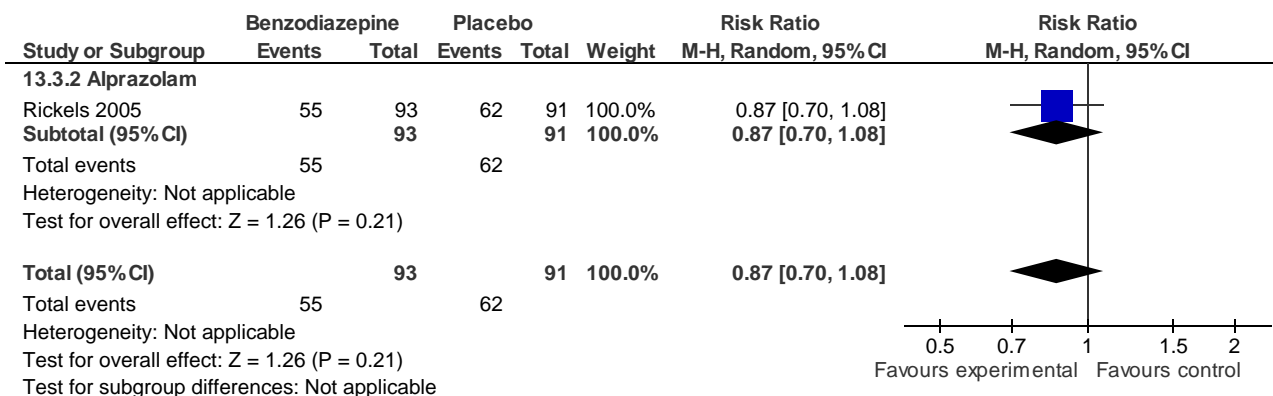


### 13 Alprazolam versus Placebo

#### 13.2 Discontinuation due to adverse events

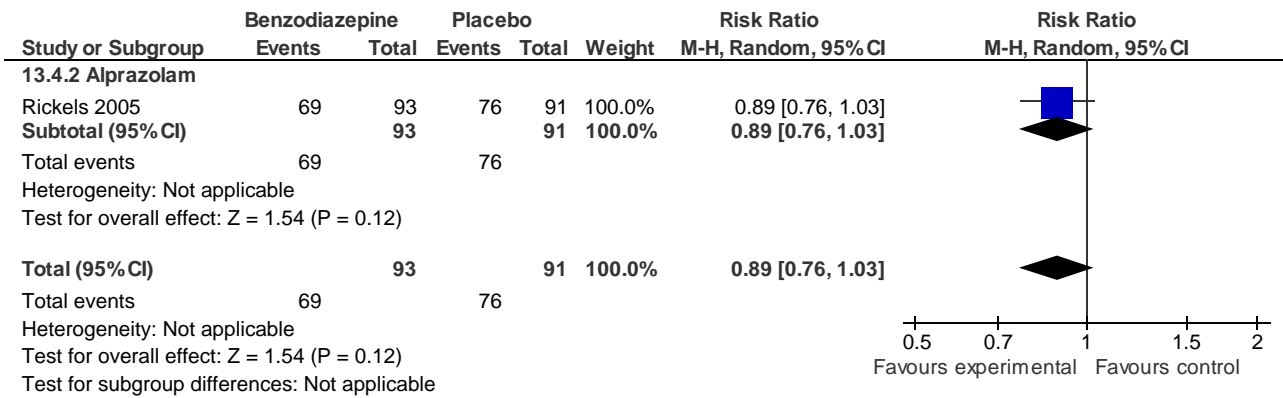


#### 13.3 Non-response

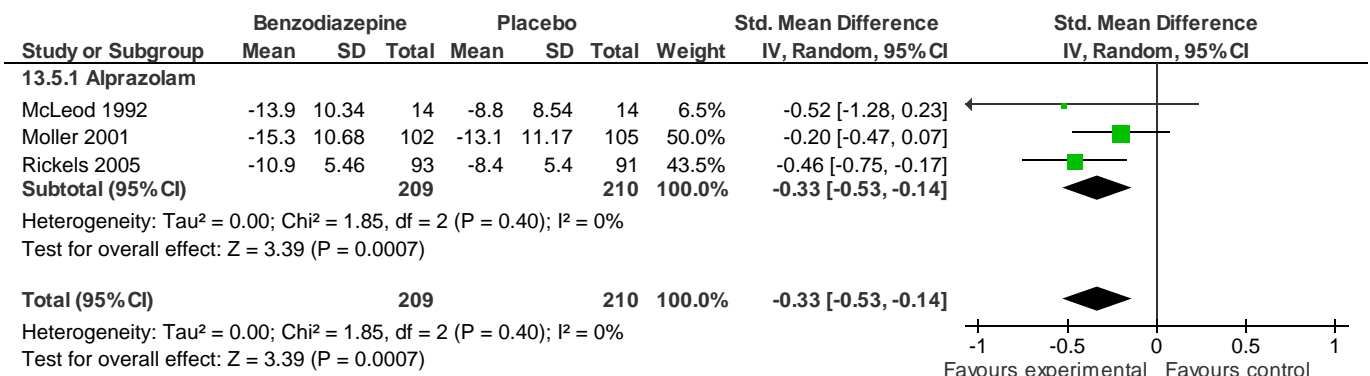


## 2 Benzodiazepine versus Placebo for GAD

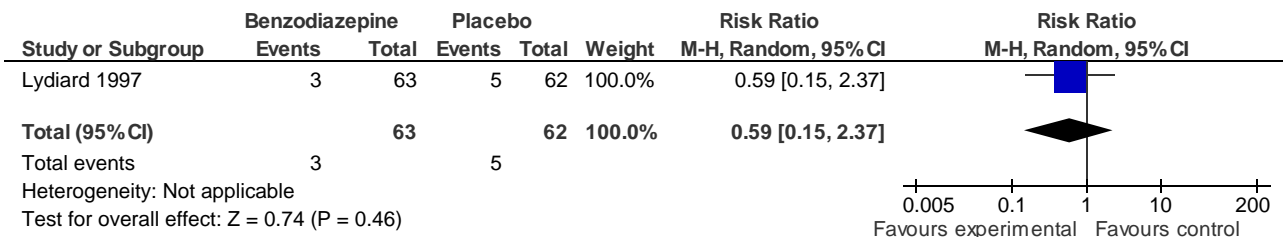
### 13.4 Non-remission



### 13.5 HAM-A

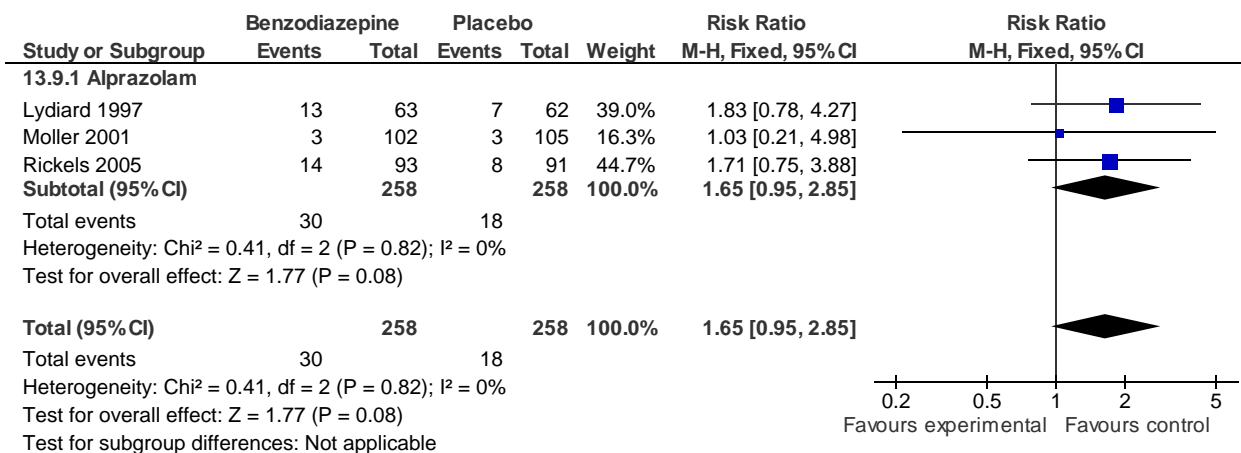


### 13.7 Insomnia

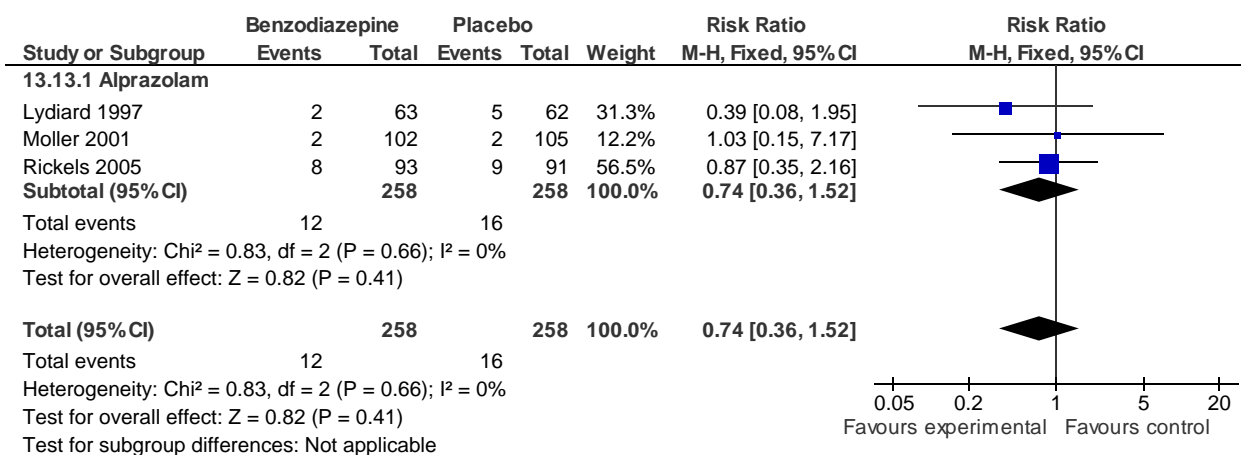


## 2 Benzodiazepine versus Placebo for GAD

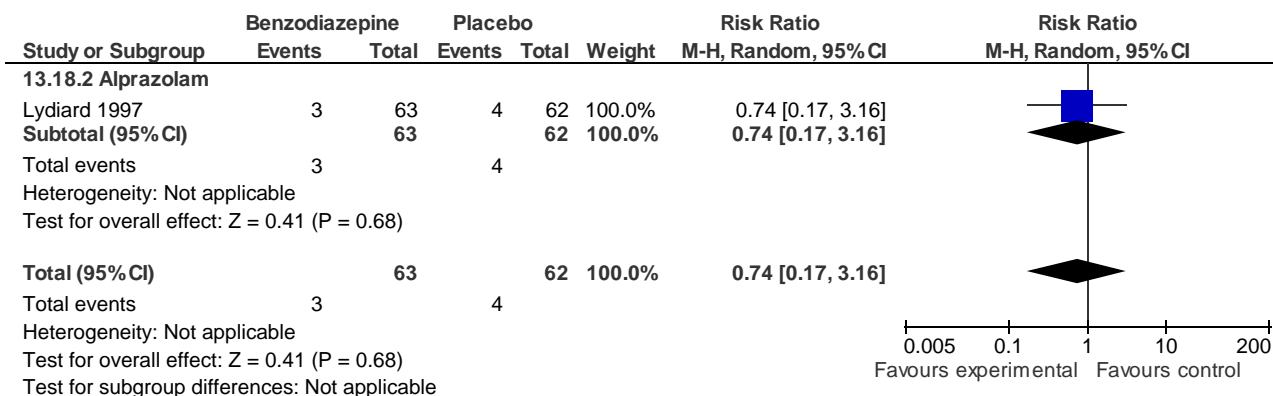
### 13.9 Dizziness



### 13.13 Nausea



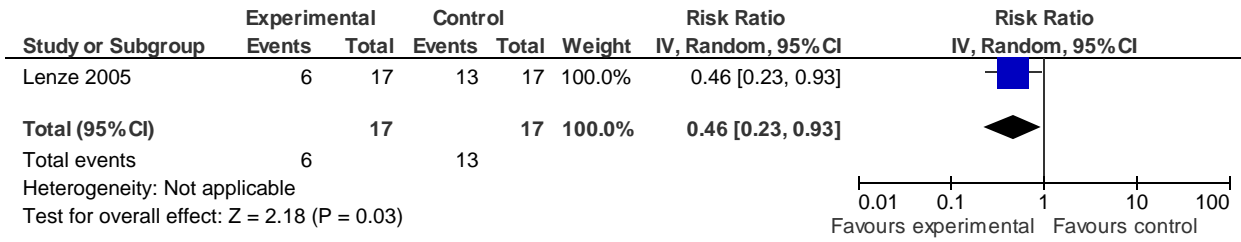
### 13.18 Fatigue



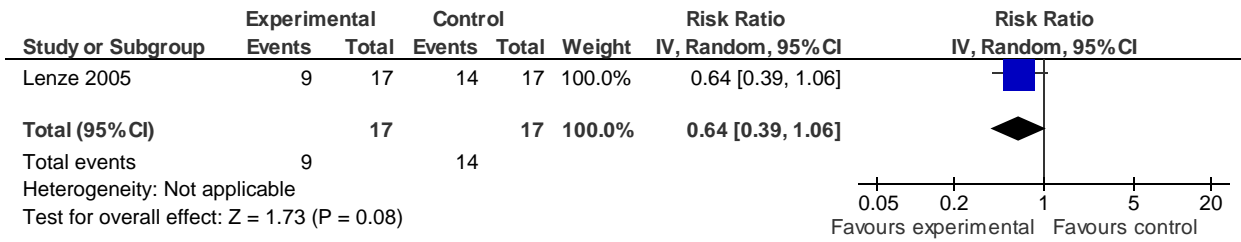
### 35 Citalopram versus Placebo

## 2 Benzodiazepine versus Placebo for GAD

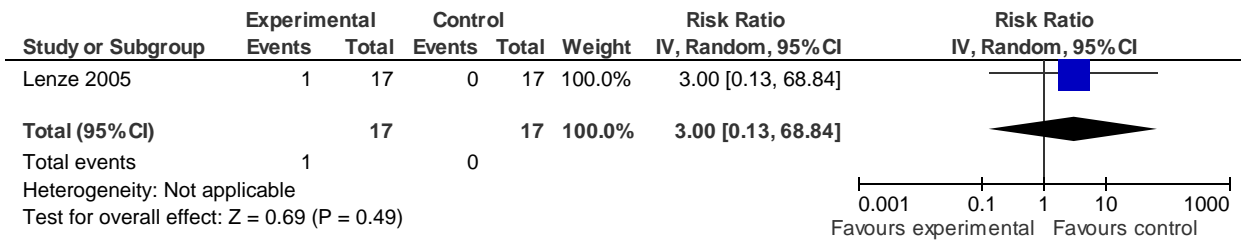
### 35.1 Non-response



### 35.2 Non-remission



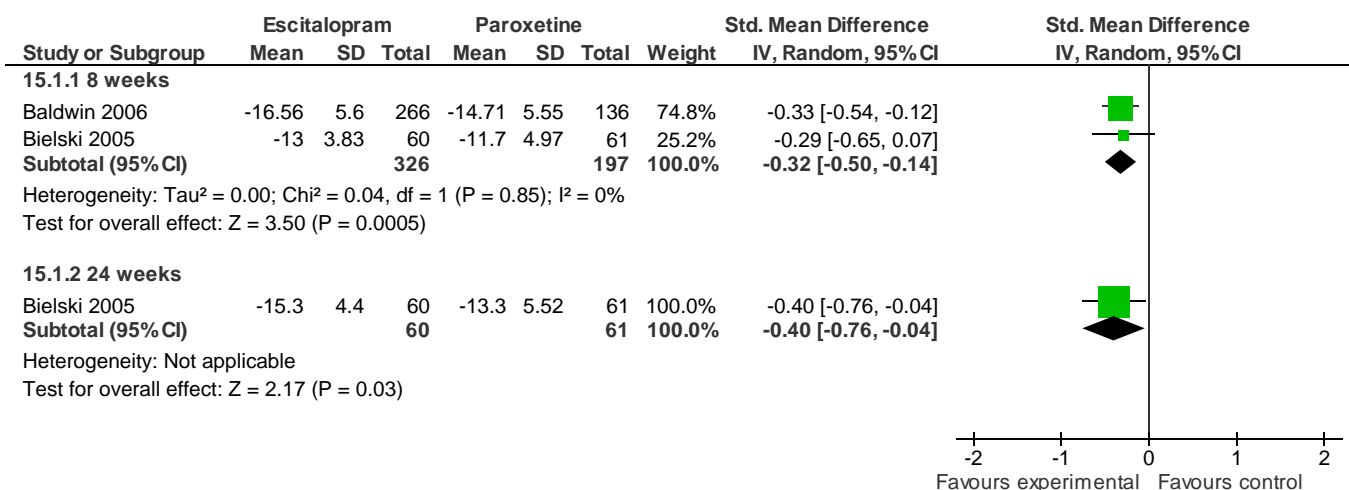
### 35.3 Discontinuation due to adverse events



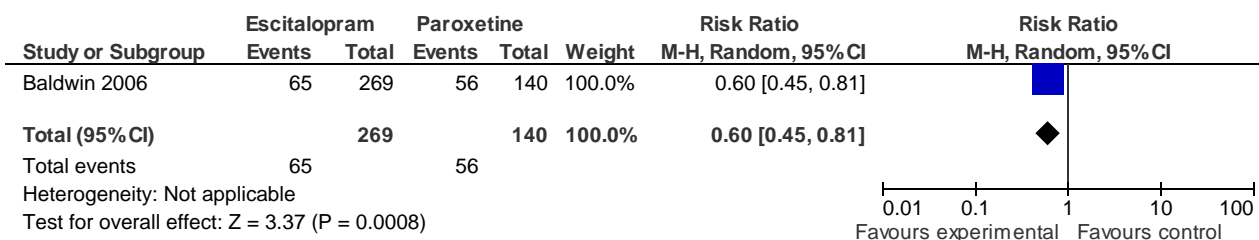
### 3 Head-to-Head trials of Pharmacological Interventions

#### 15 Escitalopram versus Paroxetine

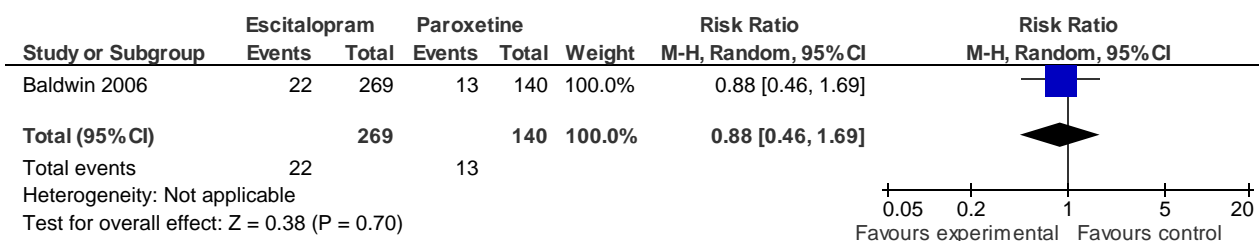
##### 15.1 HAM-A change from baseline



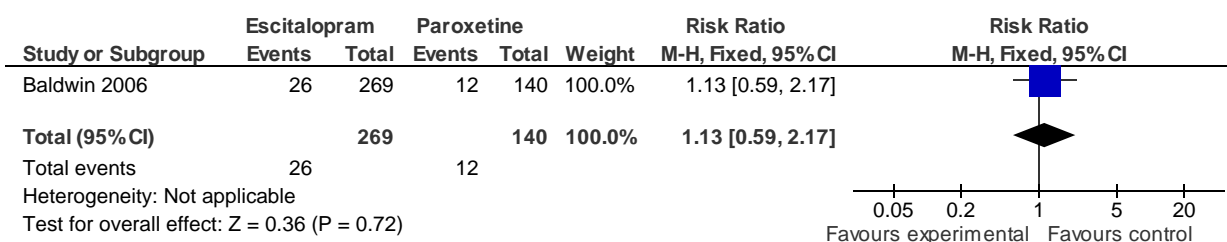
##### 15.2 Non-response



##### 15.7 Discontinuation due to adverse events



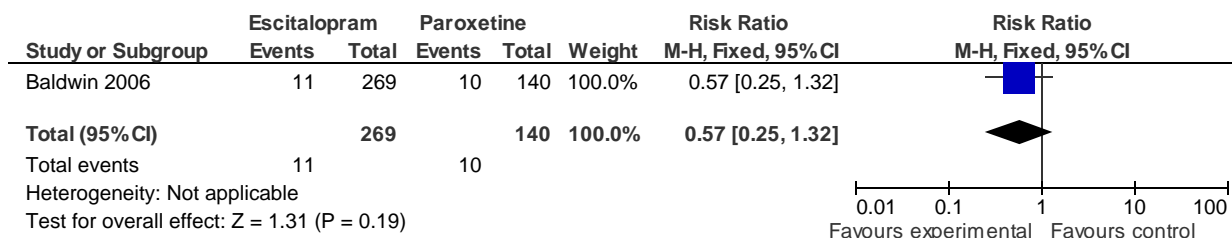
##### 15.8 Diarrhoea



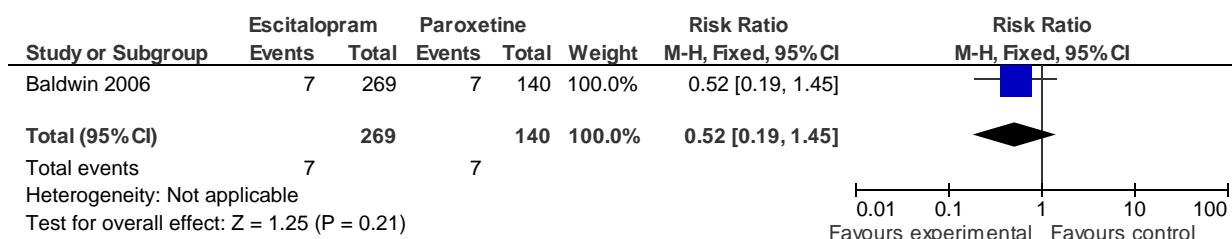


### 3 Head-to-Head trials of Pharmacological Interventions

#### 15.14 Decreased libido

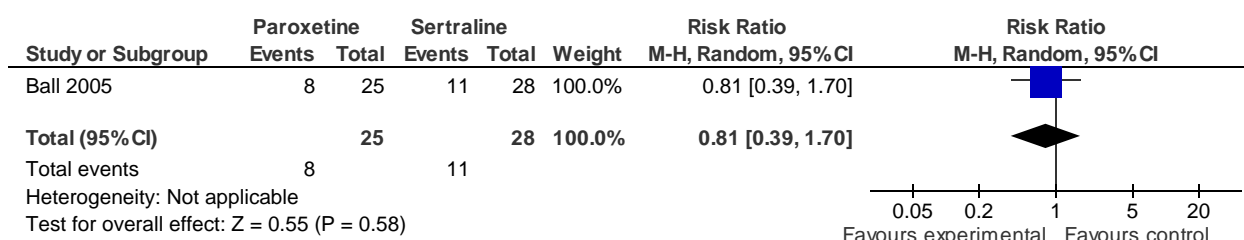


#### 15.17 Anxiety

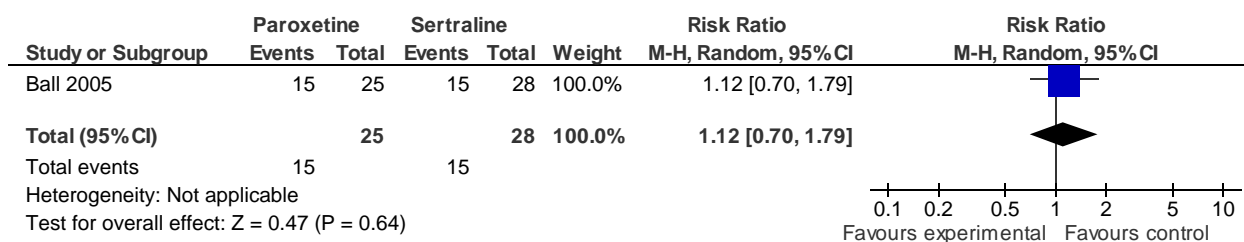


#### 16 Sertraline versus Paroxetine

##### 16.1 Non-response

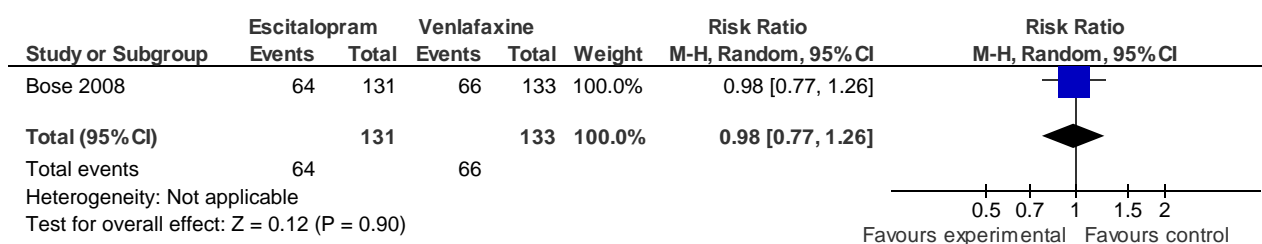


##### 16.2 Non-remission



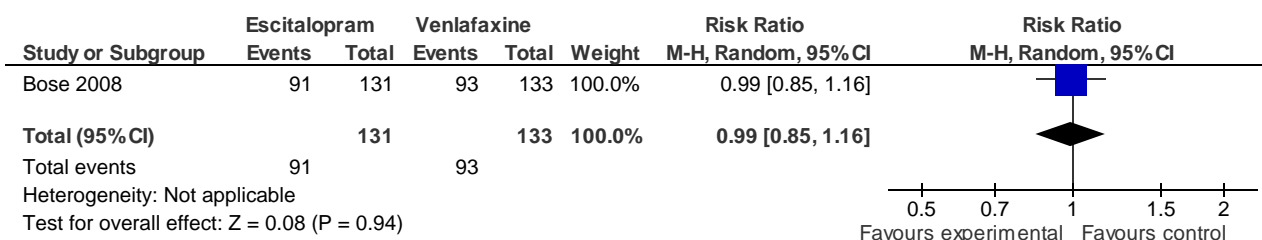
#### 17 Escitalopram versus Venlafaxine

##### 17.2 Non-response

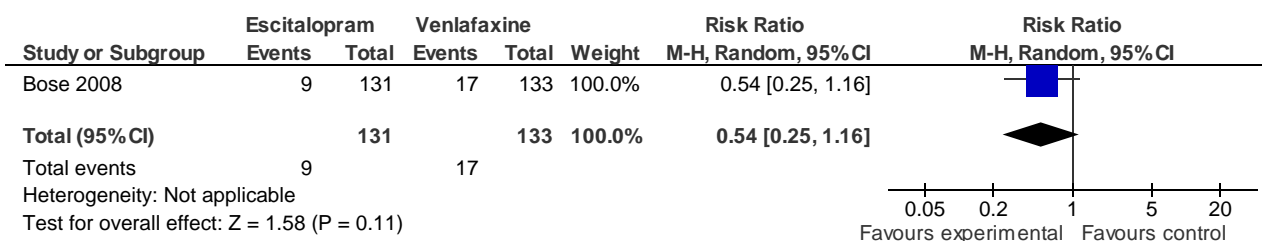


### 3 Head-to-Head trials of Pharmacological Interventions

#### 17.3 Non-remission

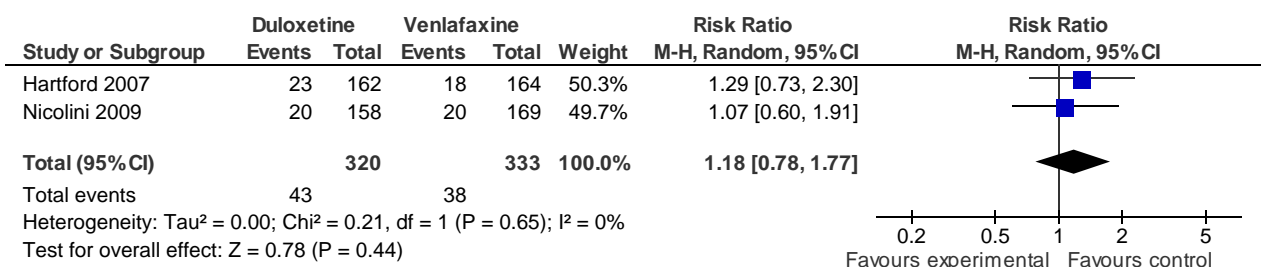


#### 17.4 Discontinuation due to adverse events

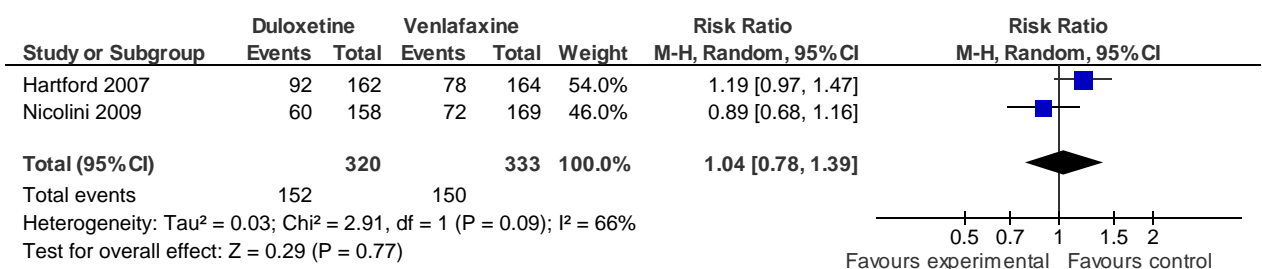


#### 21 Duloxetine versus Venlafaxine

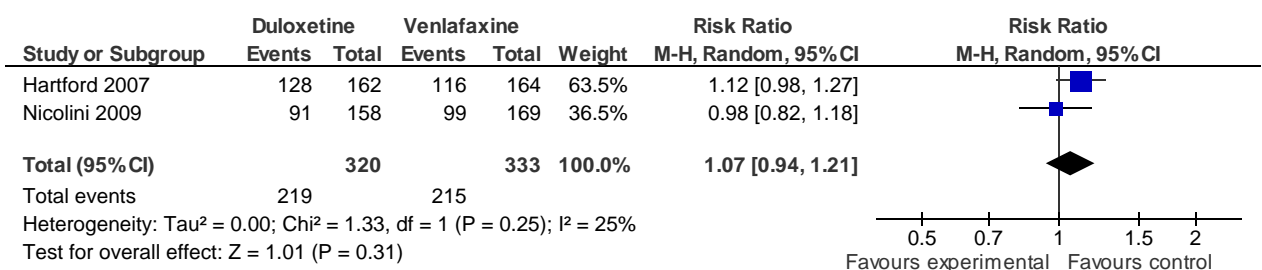
##### 21.2 Discontinuation due to adverse events



##### 21.3 Non-response

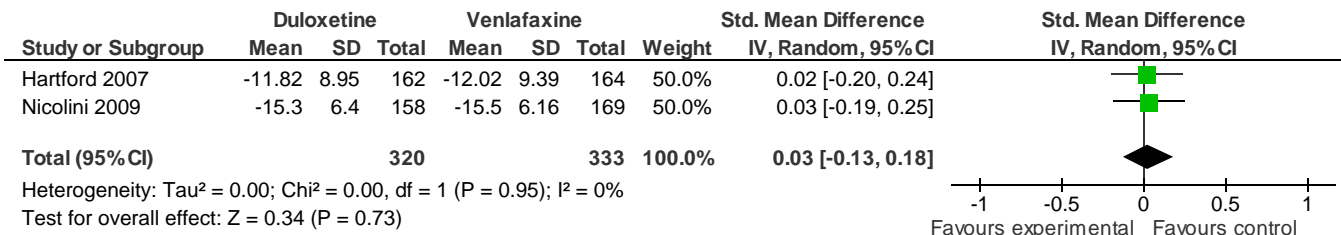


##### 21.4 Non-remission

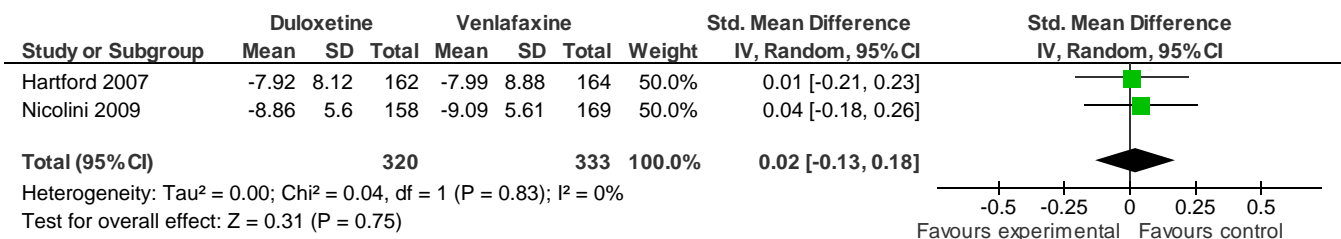


### 3 Head-to-Head trials of Pharmacological Interventions

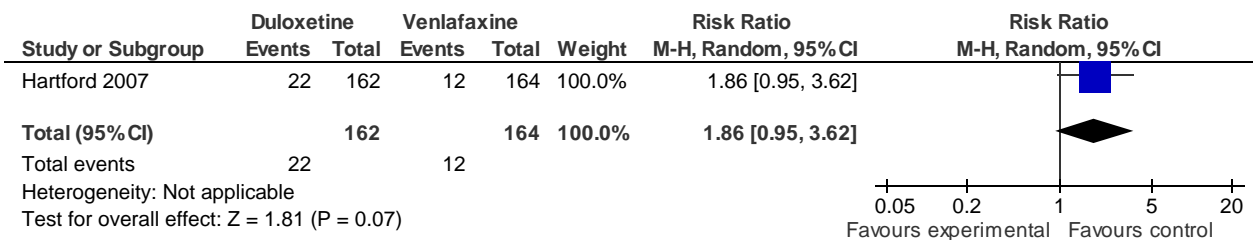
#### 21.5 HAM-A



#### 21.6 Sheehan Disability Scale



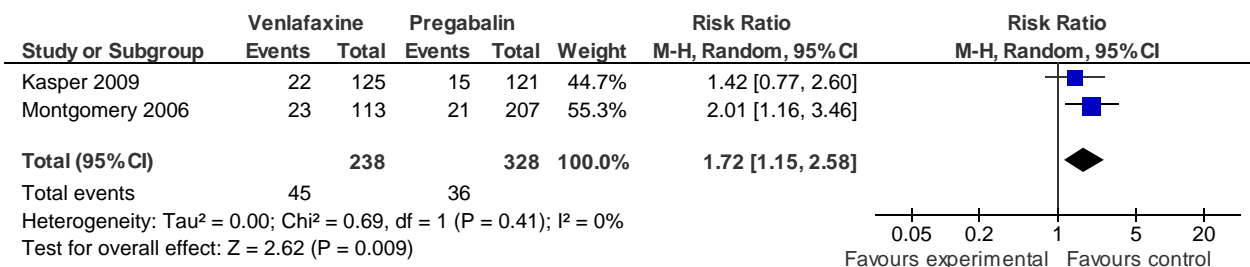
#### 21.9 Diarrhoea



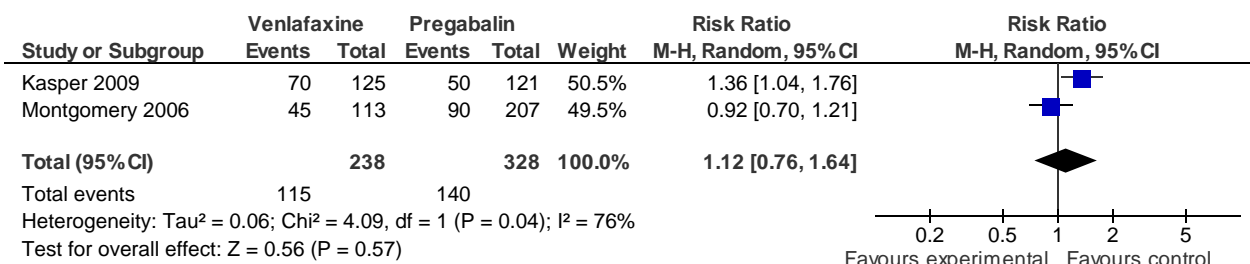
## 4 Antidepressants versus Other Pharmacological Interventions

### 14 Venlafaxine versus Pregabalin

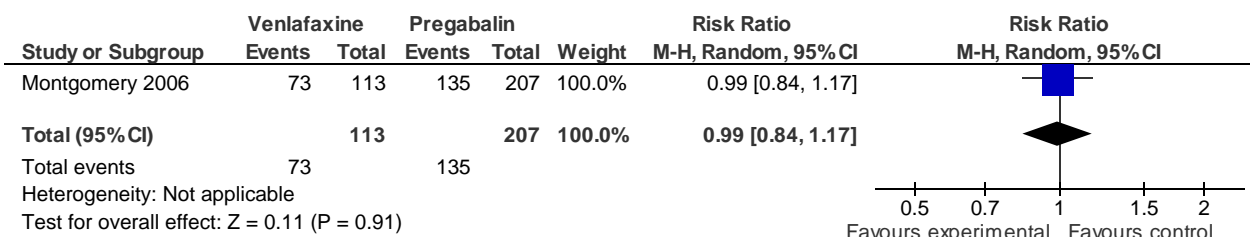
#### 14.2 Discontinuation due to adverse events



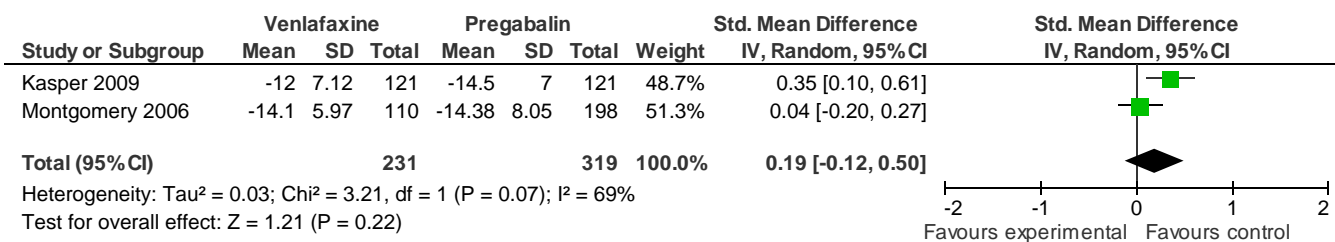
#### 14.3 Non-response



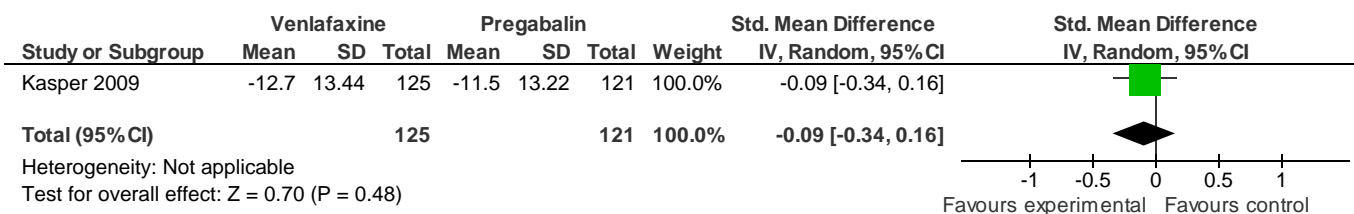
#### 14.4 Non-remission



#### 14.5 HAM-A

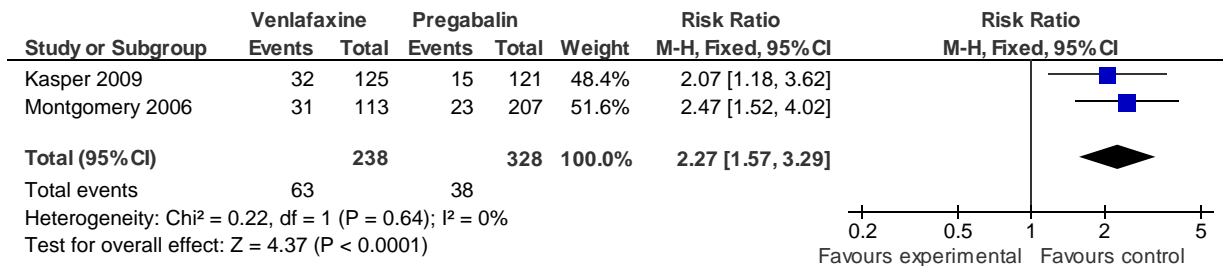


#### 14.7 Q-LES-Q

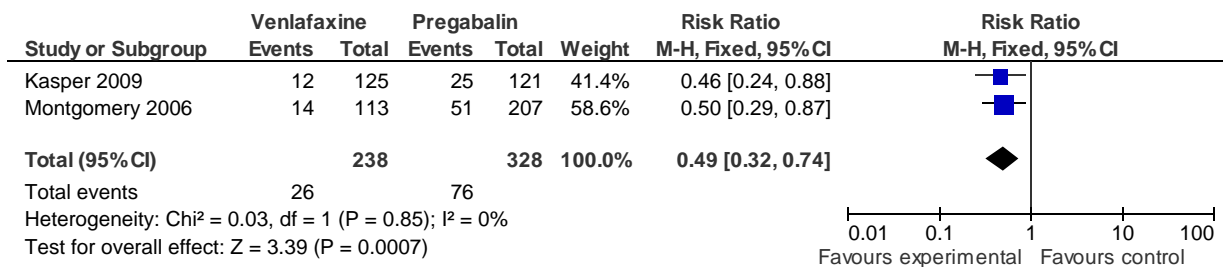


## 4 Antidepressants versus Other Pharmacological Interventions

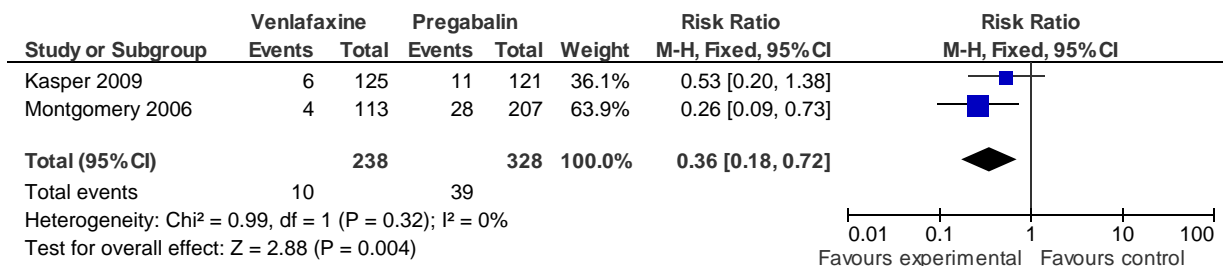
### 14.9 Nausea



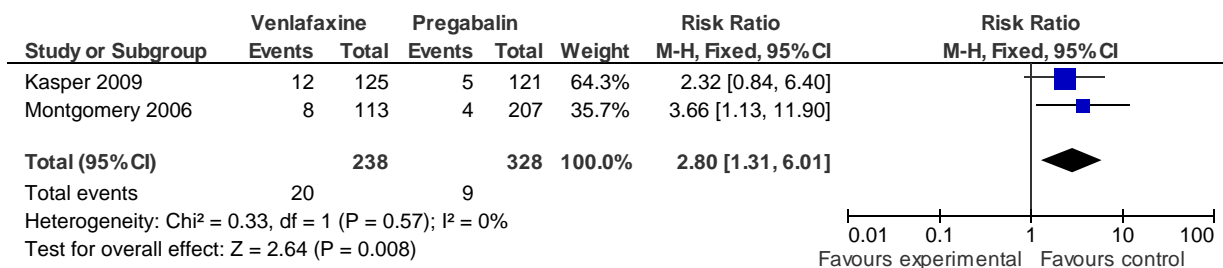
### 14.11 Dizziness



### 14.15 Somnolence

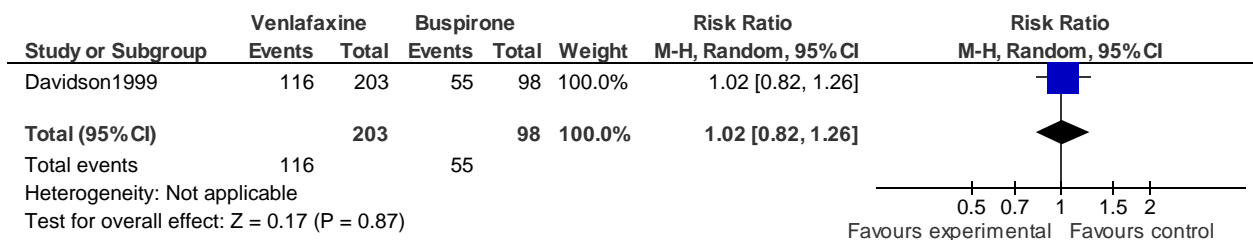


### 14.16 Insomnia



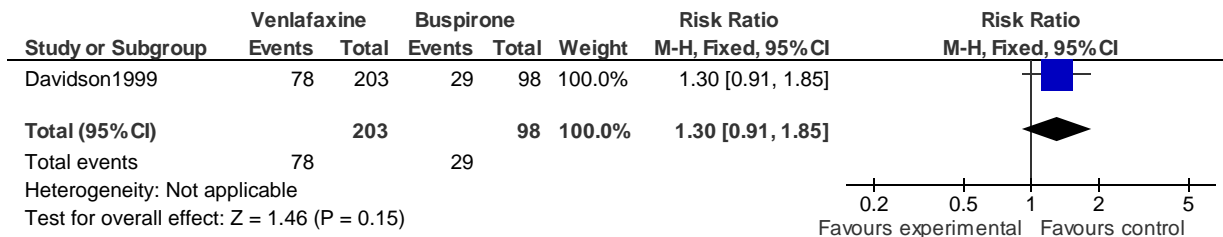
## 18 Venlafaxine versus Buspirone

### 18.1 Non-response

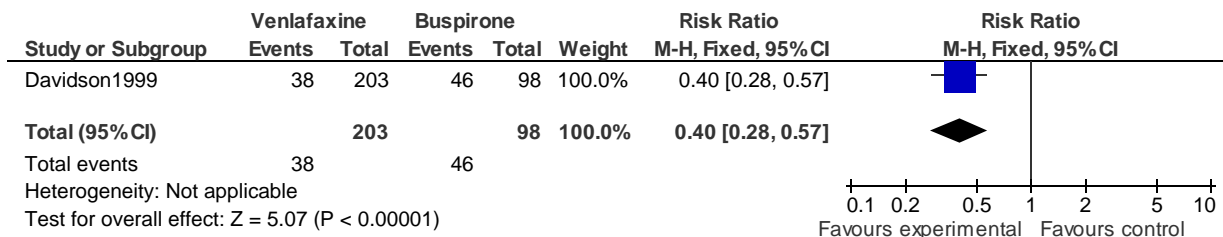


## 4 Antidepressants versus Other Pharmacological Interventions

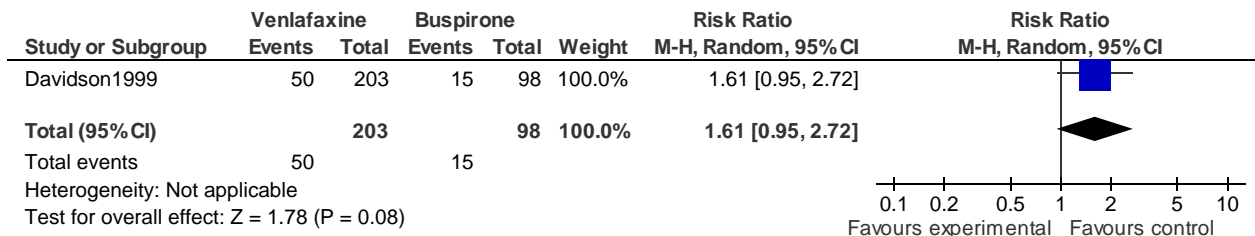
### 18.2 Nausea



### 18.3 Dizziness

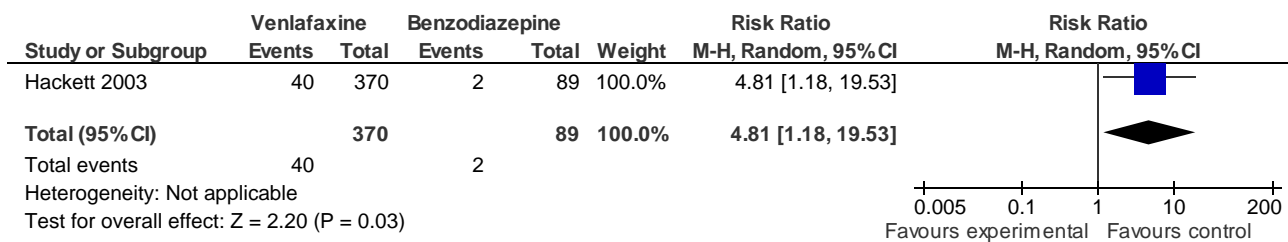


### 18.6 Discontinuation due to adverse events

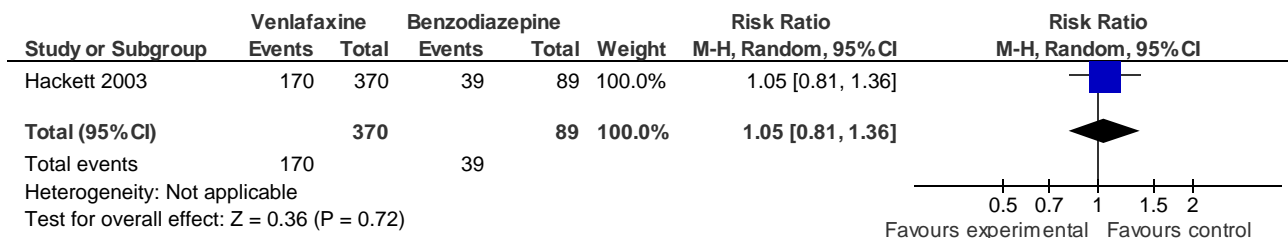


## 19 Venlafaxine versus Diazepam

### 19.2 Discontinuation due to adverse events



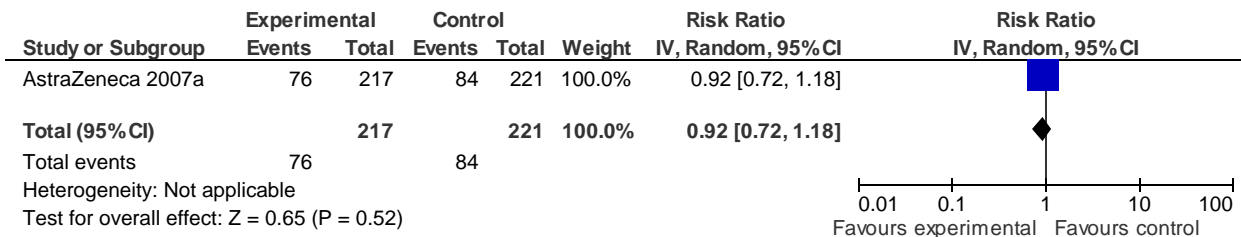
### 19.3 Non-response



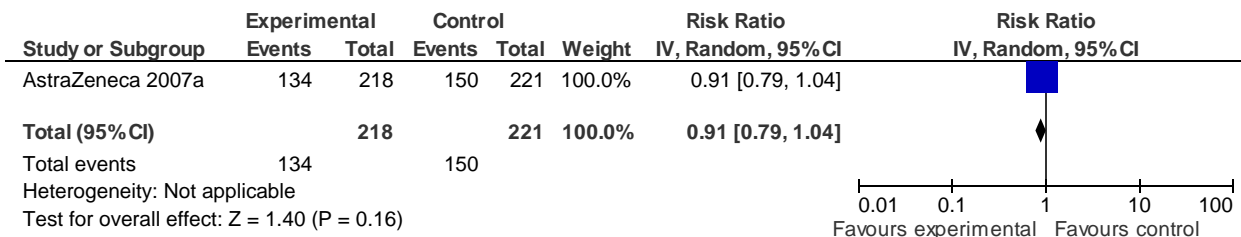
## 23 Quetiapine 50mg versus Paroxetine

## 4 Antidepressants versus Other Pharmacological Interventions

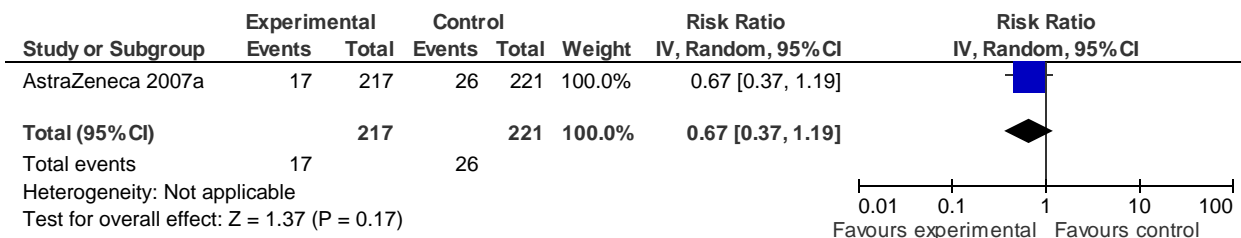
### 23.1 Non-response



### 23.2 Non-remission

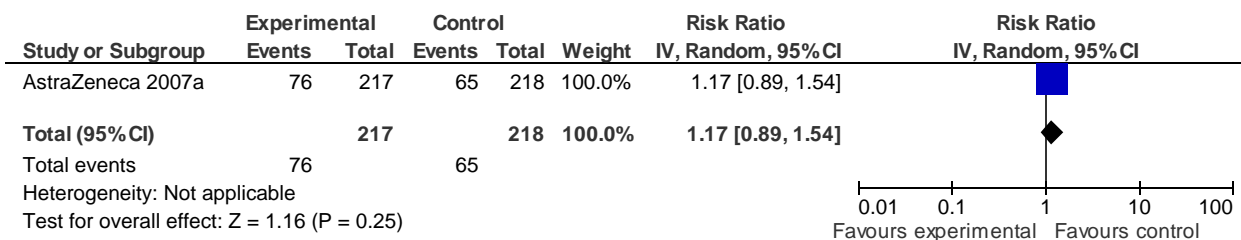


### 23.3 Discontinuation due to adverse events

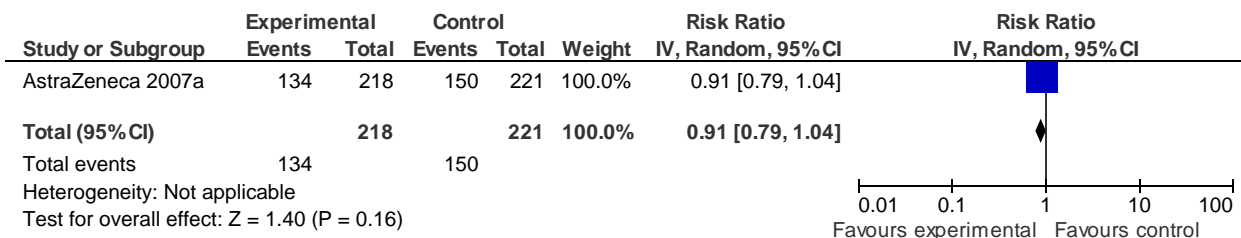


## 24 Quetiapine 150mg versus Paroxetine

### 24.1 Non-response

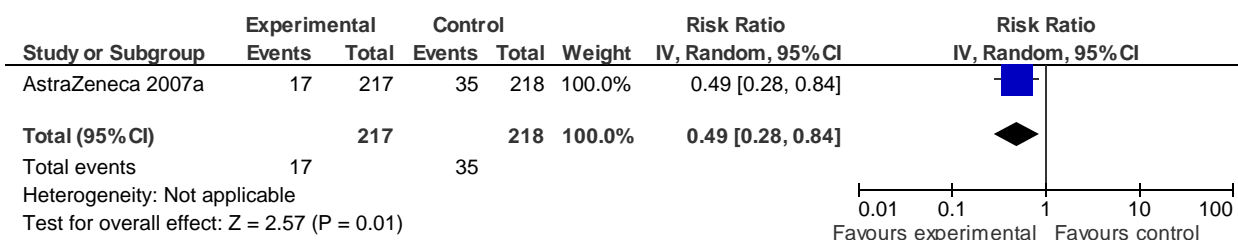


### 24.2 Non-remission



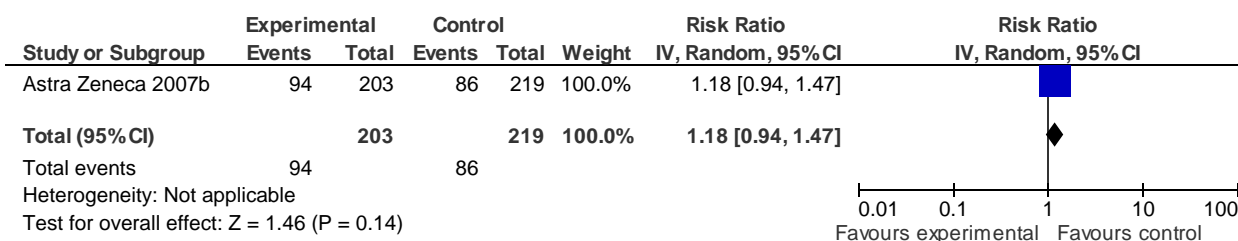
## 4 Antidepressants versus Other Pharmacological Interventions

### 24.3 Discontinuation due to adverse events

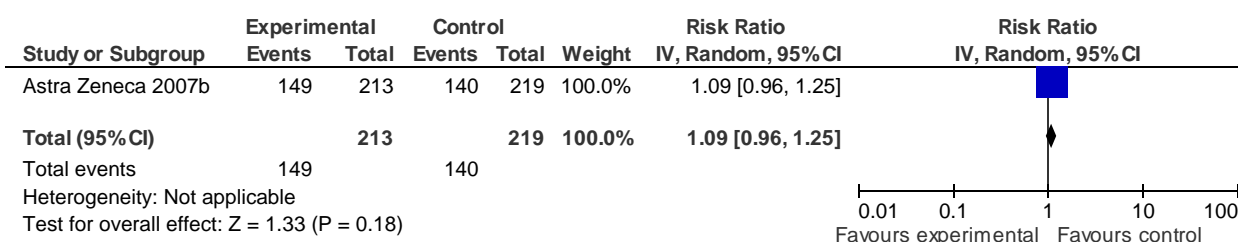


### 33 Quetiapine 150mg versus Escitalopram

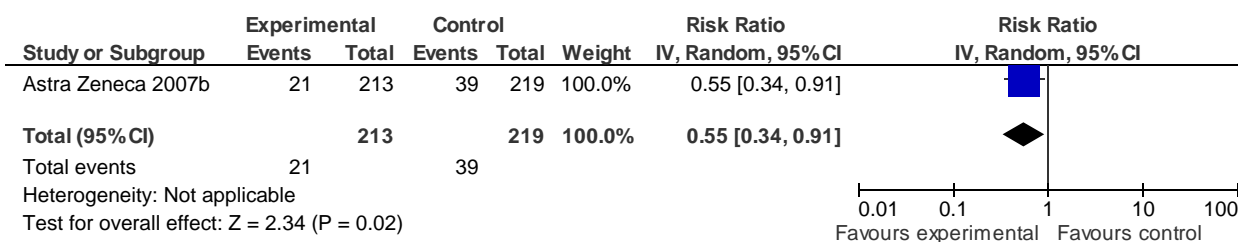
#### 33.1 Non-response



#### 33.2 Non-remission

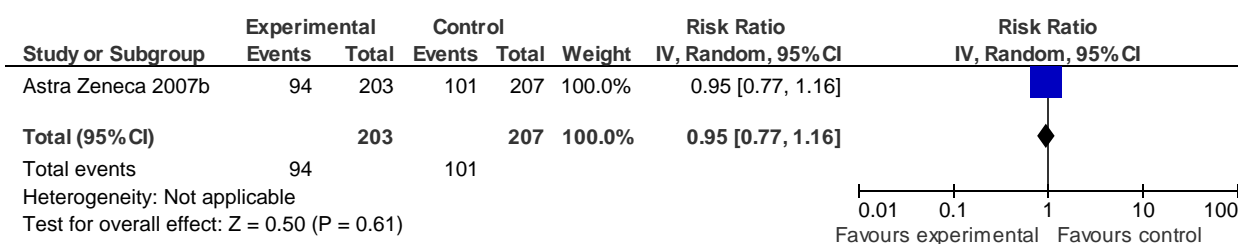


### 33.3 Discontinuation due to adverse events



### 34 Quetiapine 300mg versus Escitalopram

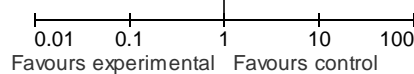
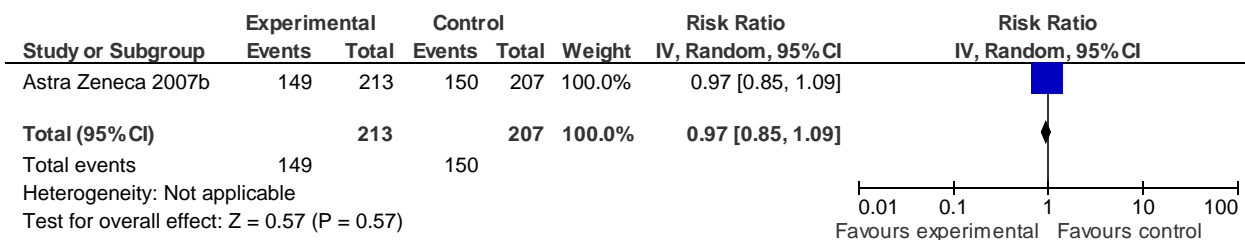
#### 34.1 Non-response



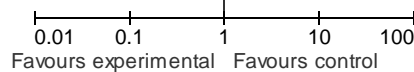
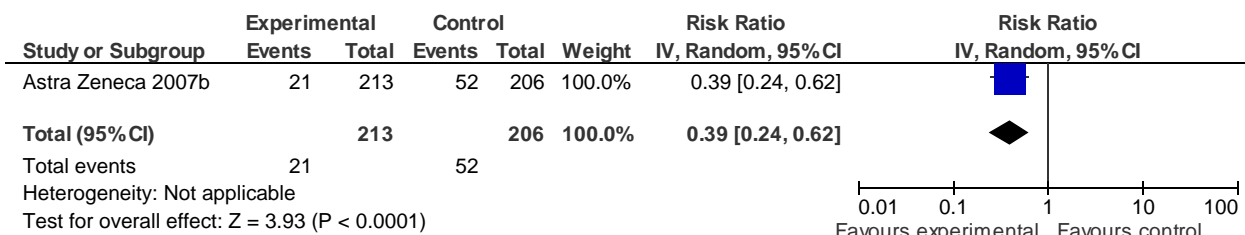


## 4 Antidepressants versus Other Pharmacological Interventions

### 34.2 Non-remission



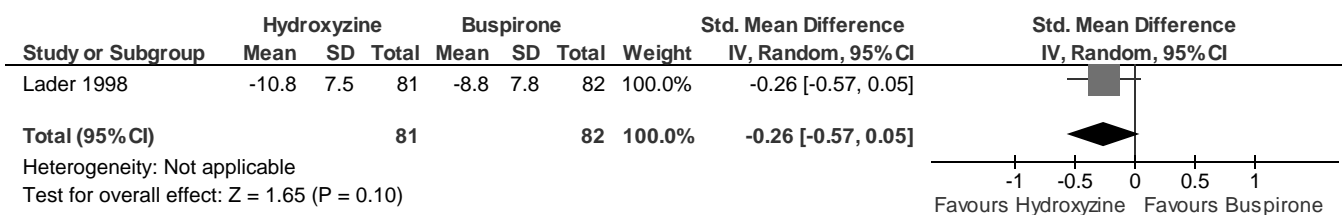
### 34.3 Discontinuation due to adverse events



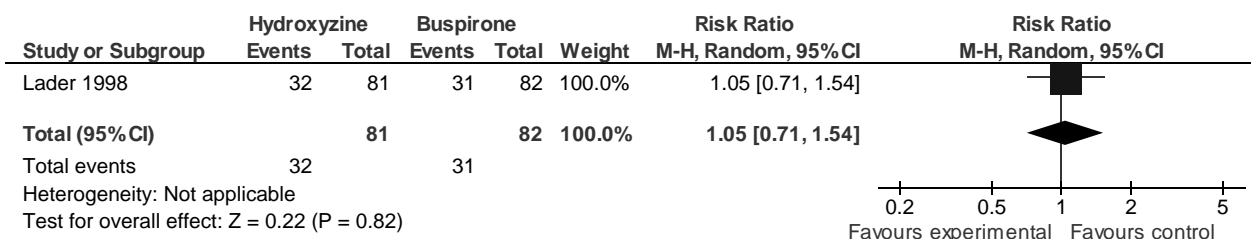
## 5 Other Pharmacological Interventions (other than antidepressants)

### 20 Hydroxyzine versus Buspirone

#### 20.2 HAM-A

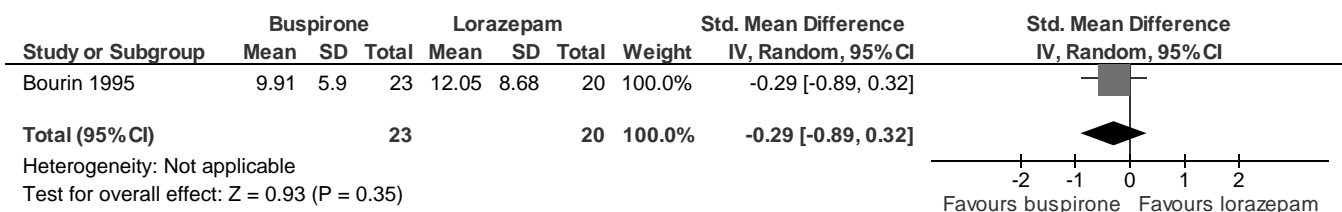


#### 20.4 At least one side effect



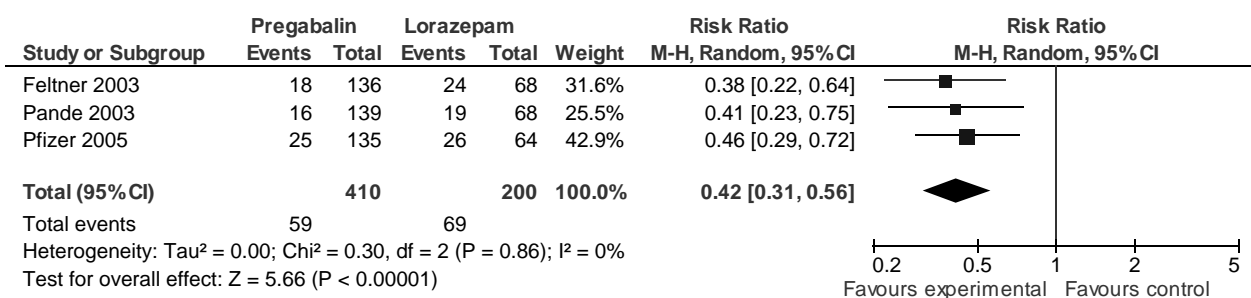
### 22 Buspirone versus Lorazepam

#### 22.1 HAM-A



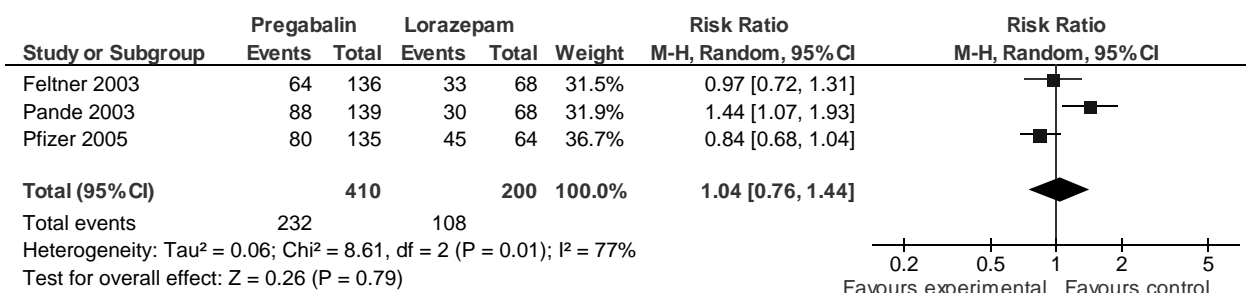
### 25 Pregabalin versus Lorazepam

#### 25.2 Discontinuation due to adverse events

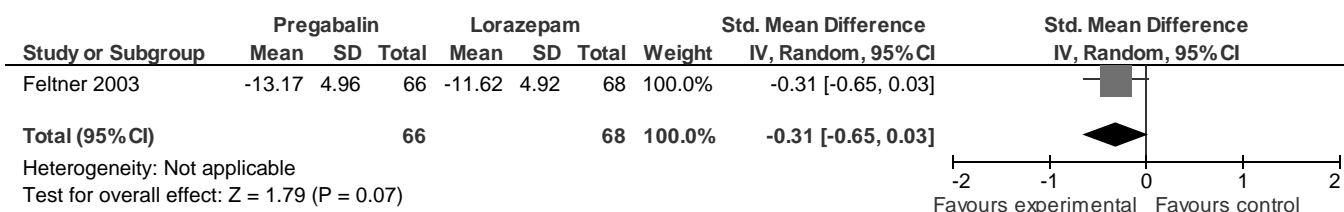


## 5 Other Pharmacological Interventions (other than antidepressants)

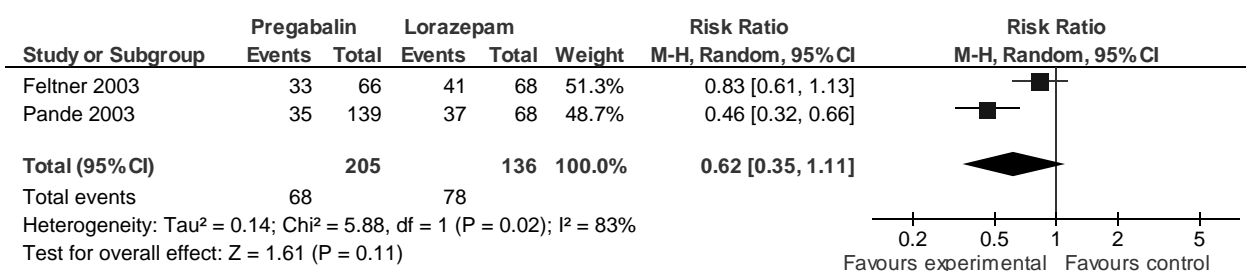
### 25.3 Non-response



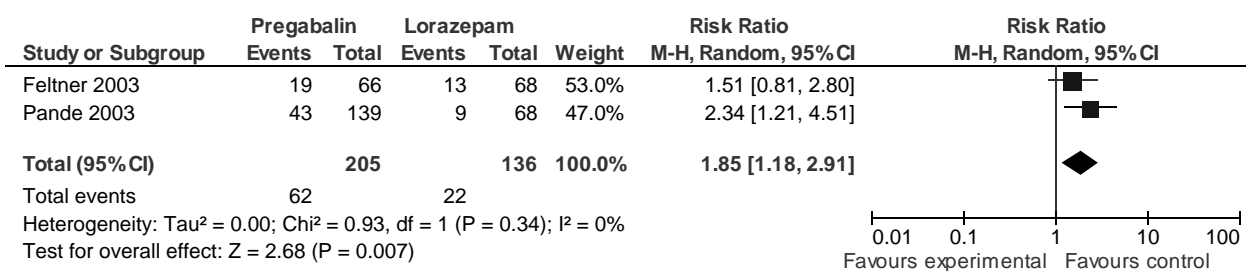
### 25.4 HAM-A



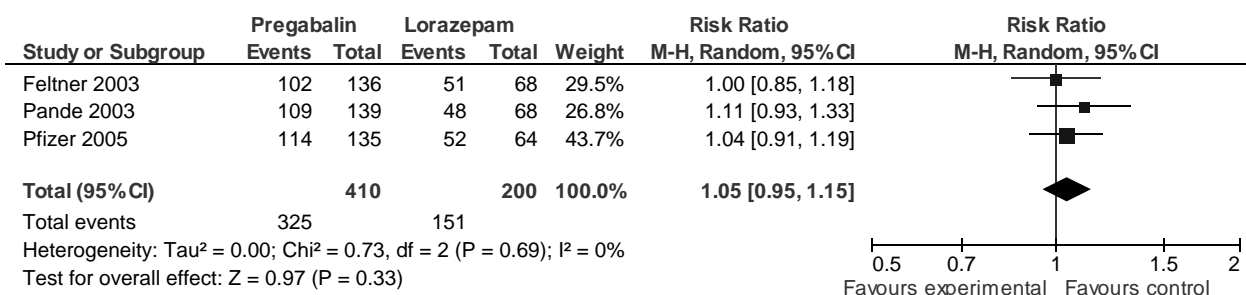
### 25.5 Somnolence



### 25.6 Dizziness



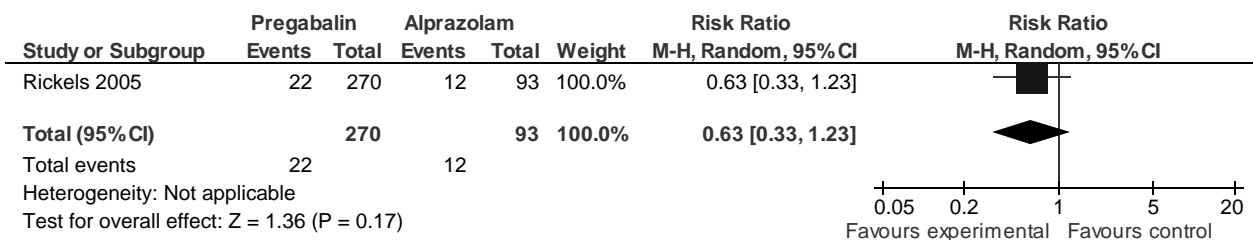
### 25.7 Non-remission



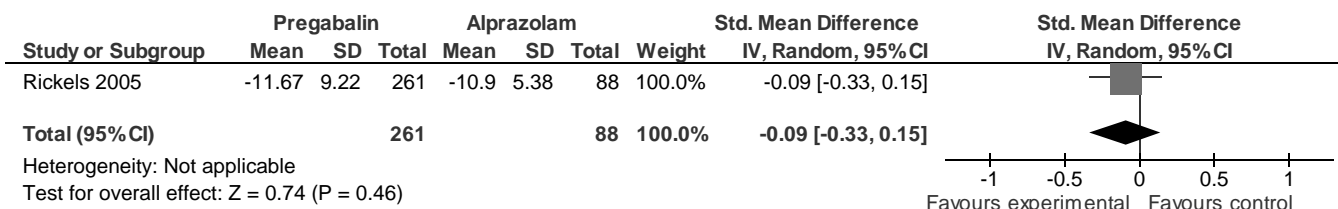
### 26 Pregabalin versus Alprazolam

## 5 Other Pharmacological Interventions (other than antidepressants)

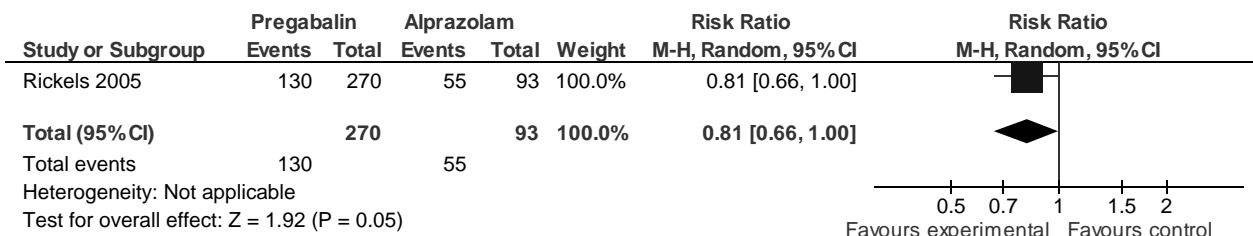
### 26.2 Discontinuation due to adverse events



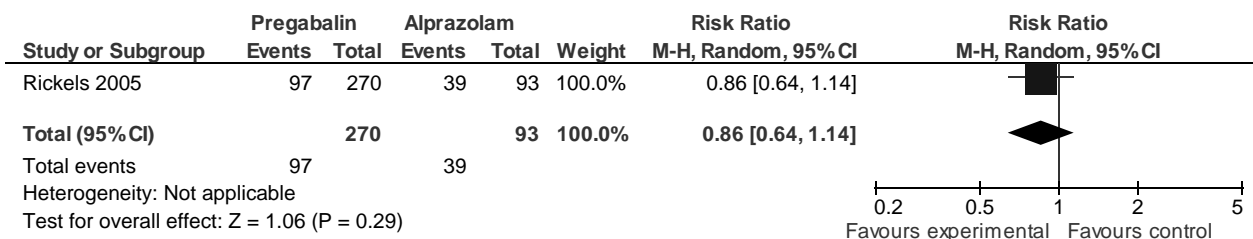
### 26.3 HAM-A



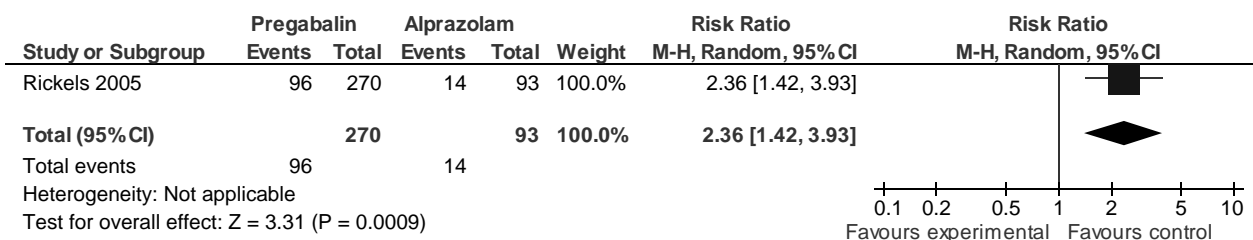
### 26.4 Non-response



### 26.5 Somnolence

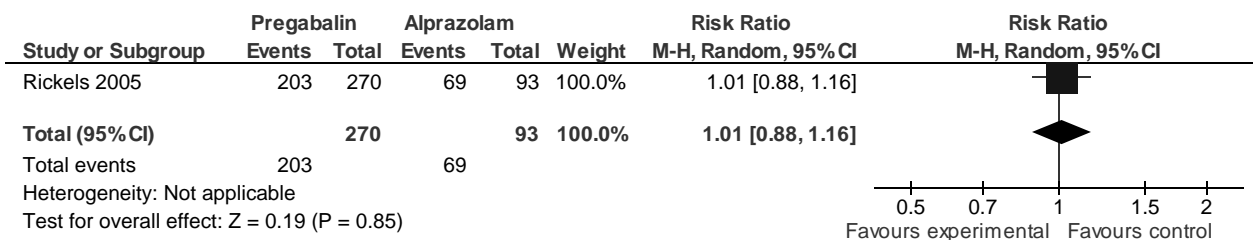


### 26.6 Dizziness



## 5 Other Pharmacological Interventions (other than antidepressants)

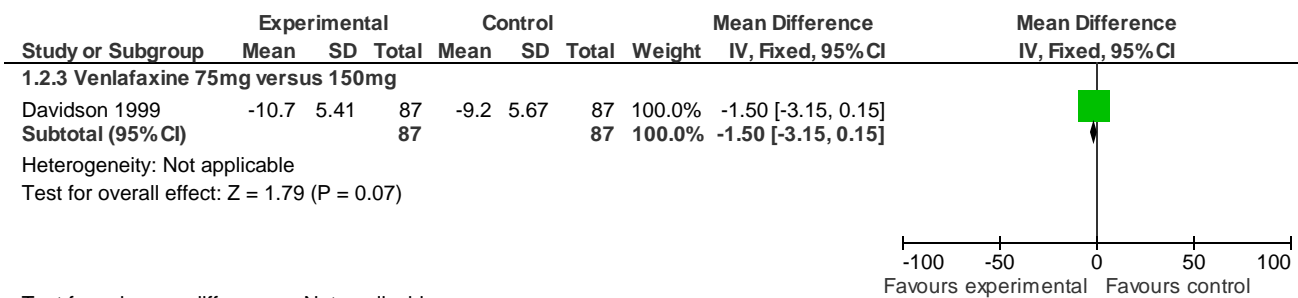
### 26.7 Non-remission



## 6 Comparing the Effectiveness of Different Dosages

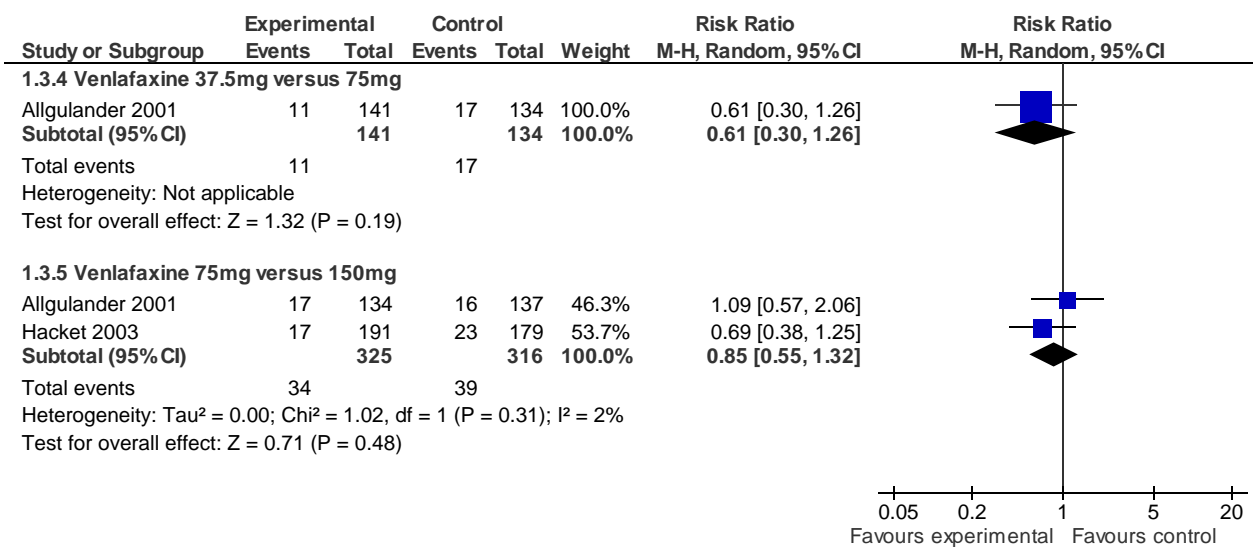
### 1 Venlafaxine

#### 1.2 HAM-A

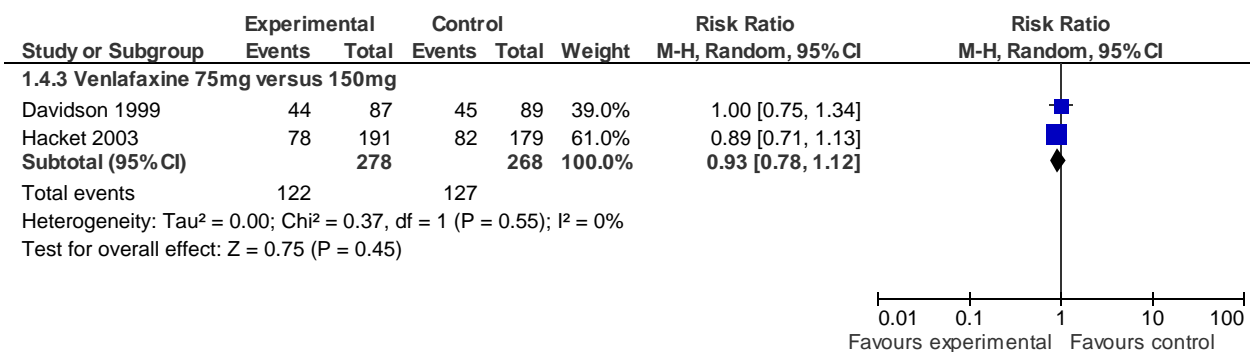


Test for subgroup differences: Not applicable

#### 1.3 Discontinuation due to Adverse Events

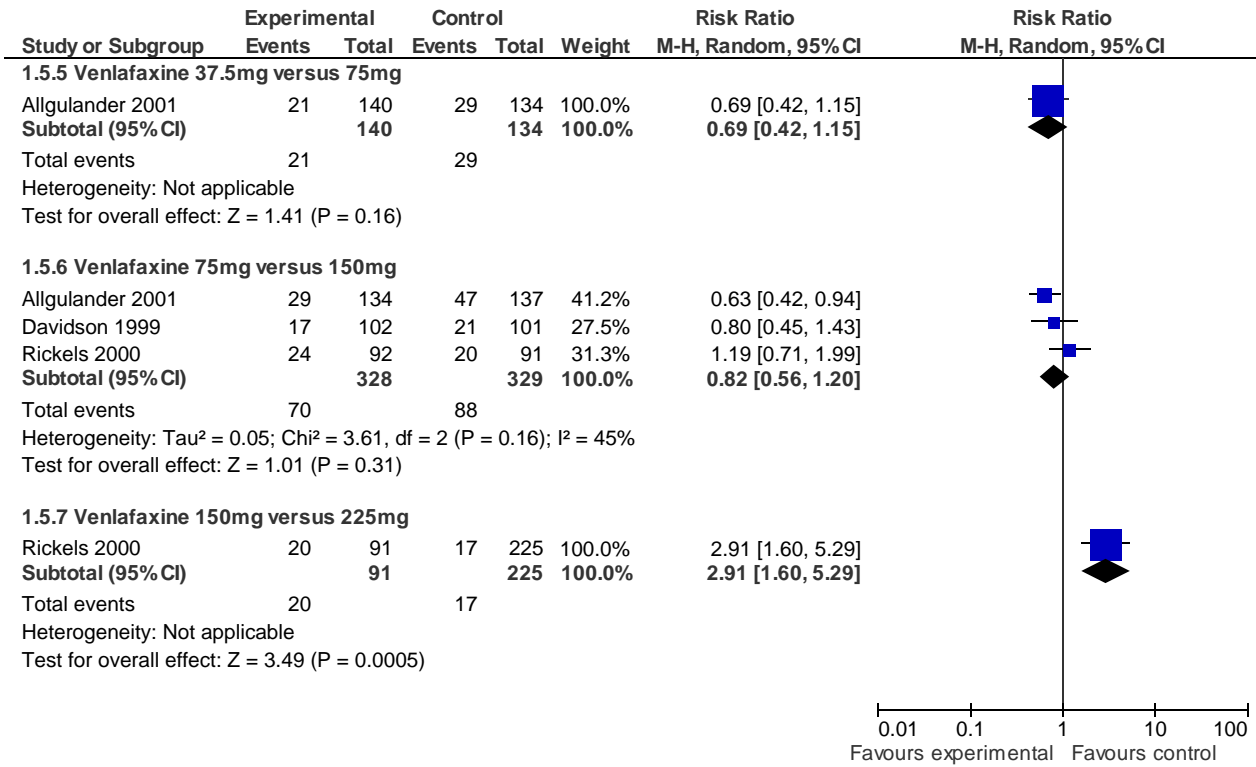


#### 1.4 Non Response



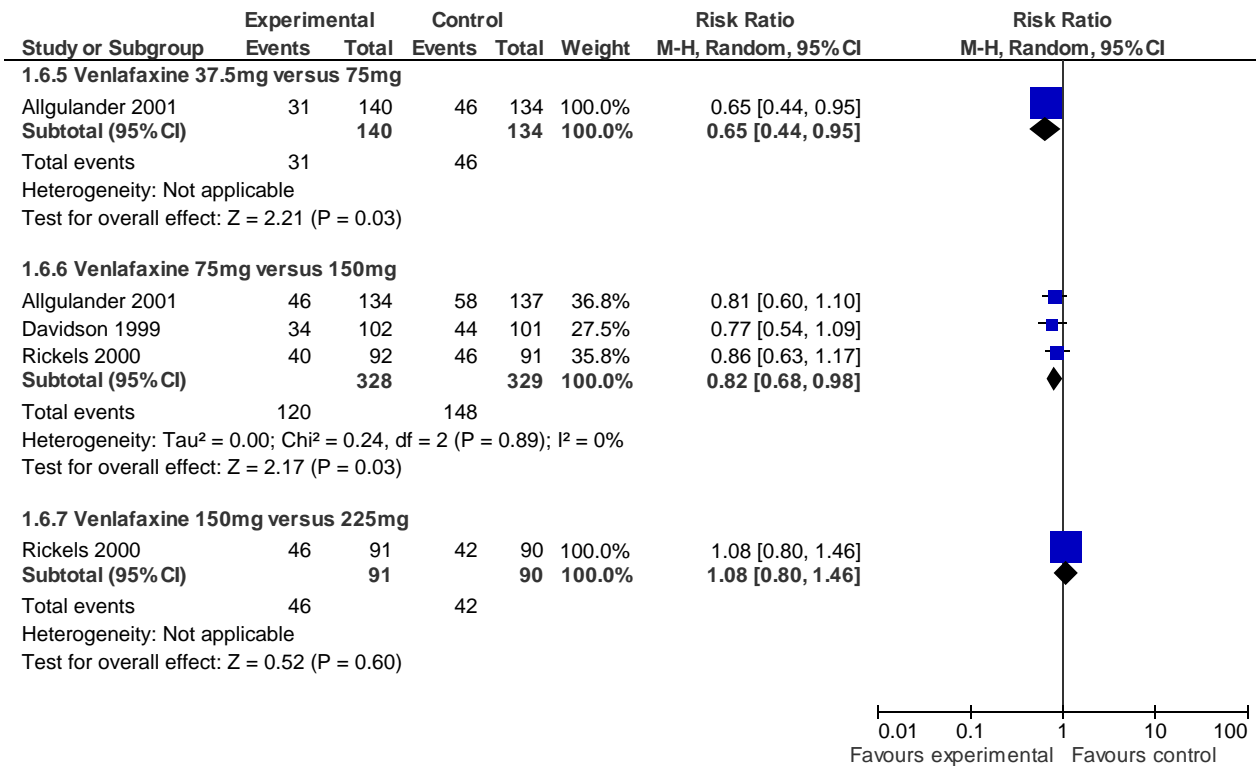
## 6 Comparing the Effectiveness of Different Dosages

### 1.5 Dizziness

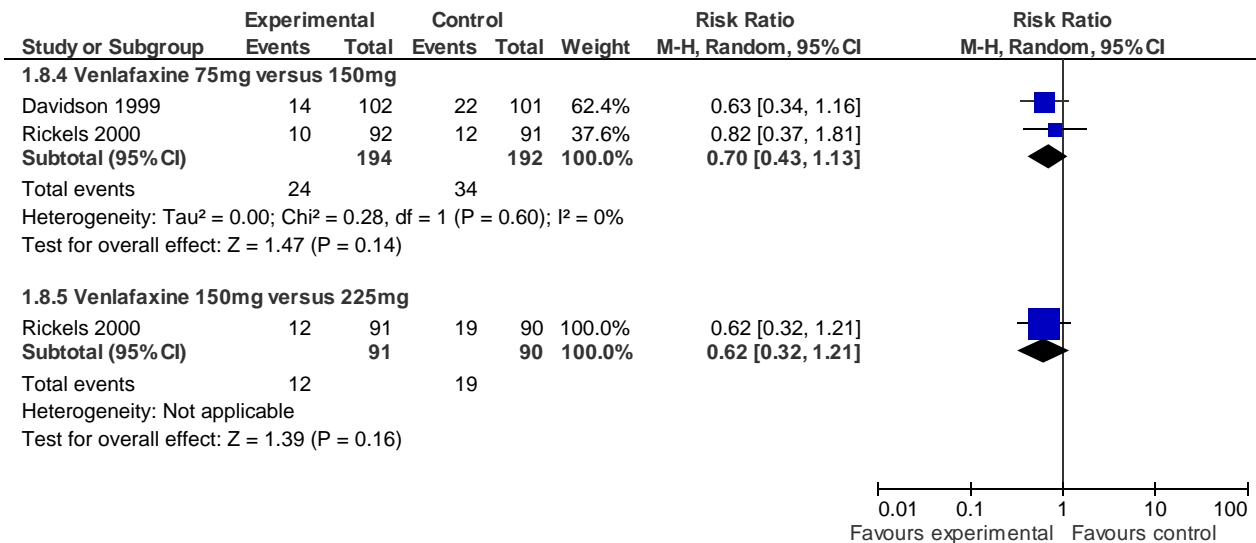


## 6 Comparing the Effectiveness of Different Dosages

### 1.6 Nausea



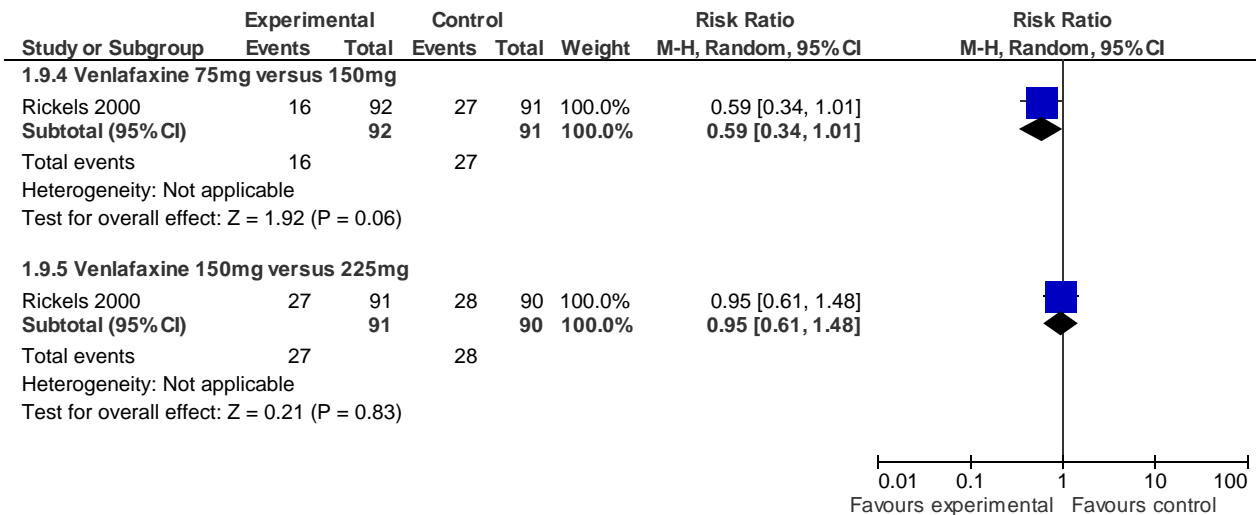
### 1.8 Asthenia



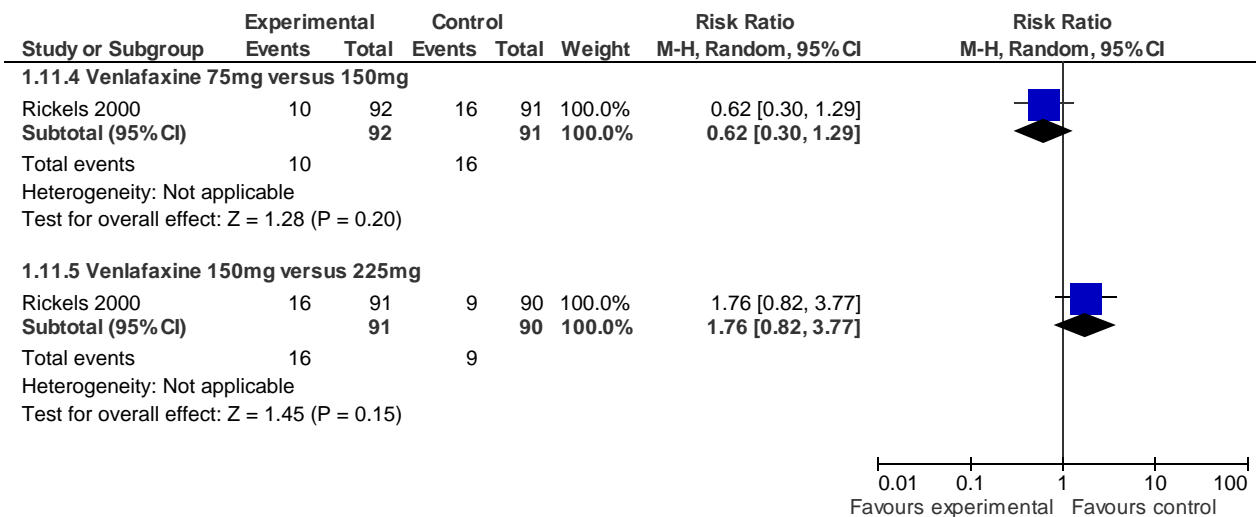


## 6 Comparing the Effectiveness of Different Dosages

### 1.9 Insomnia



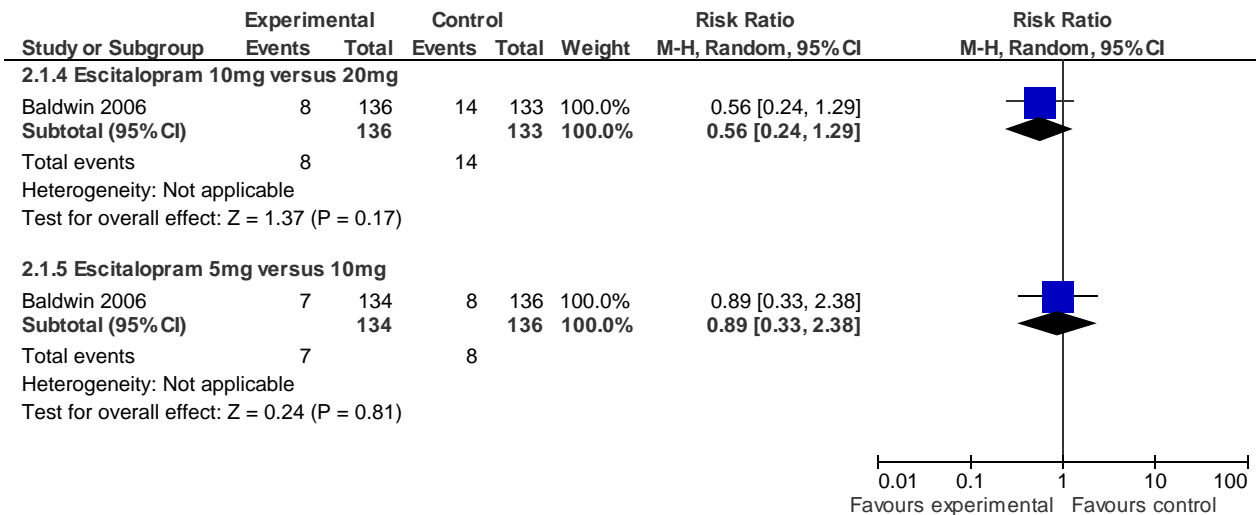
### 1.11 Nervousness



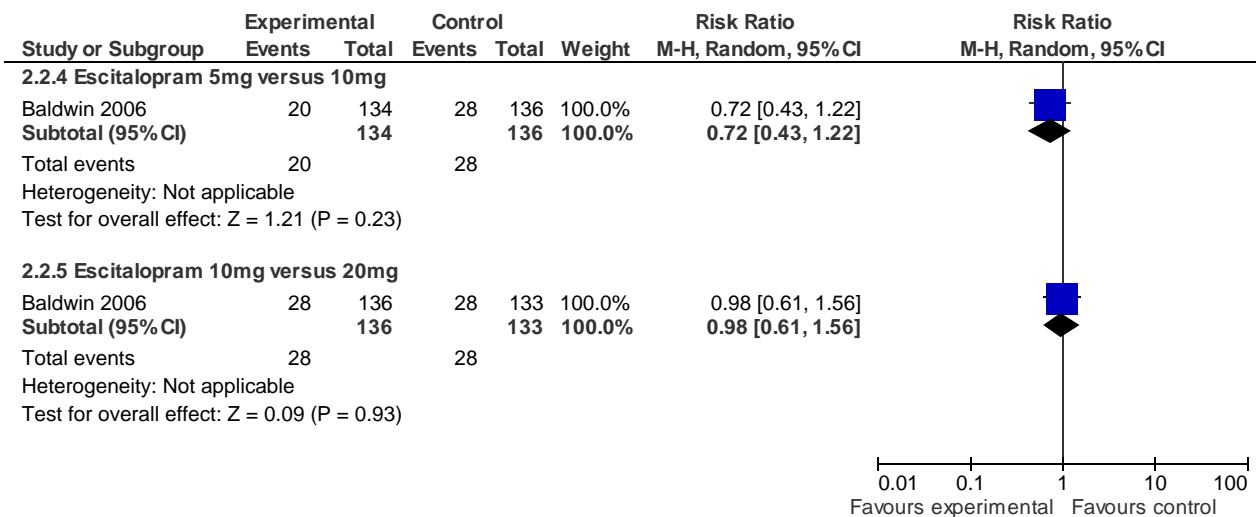
### 2 Escitalopram

## 6 Comparing the Effectiveness of Different Dosages

### 2.1 Discontinuation due to Adverse events

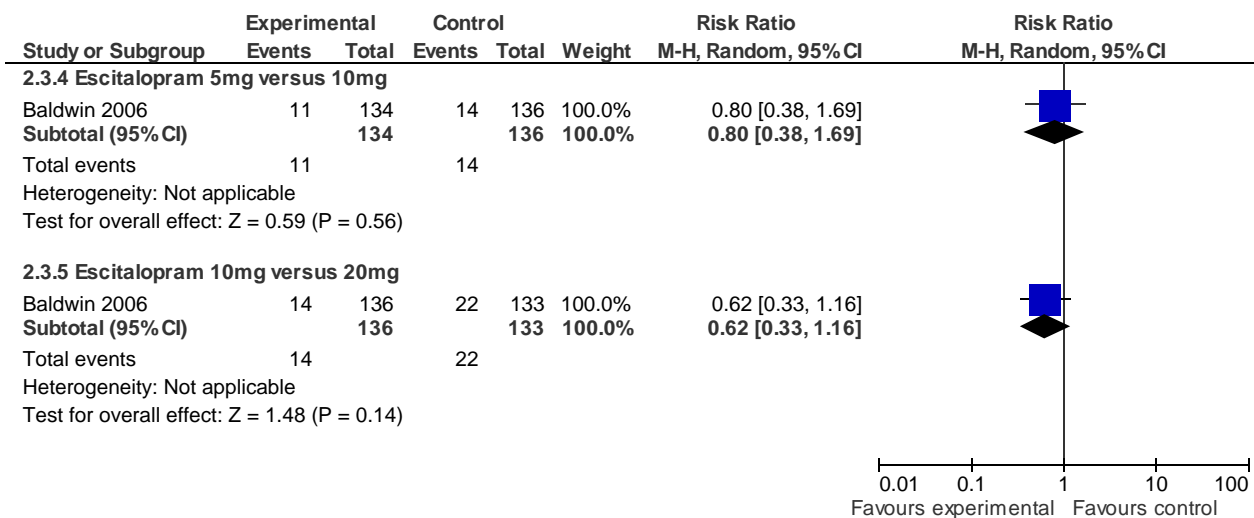


### 2.2 Nausea

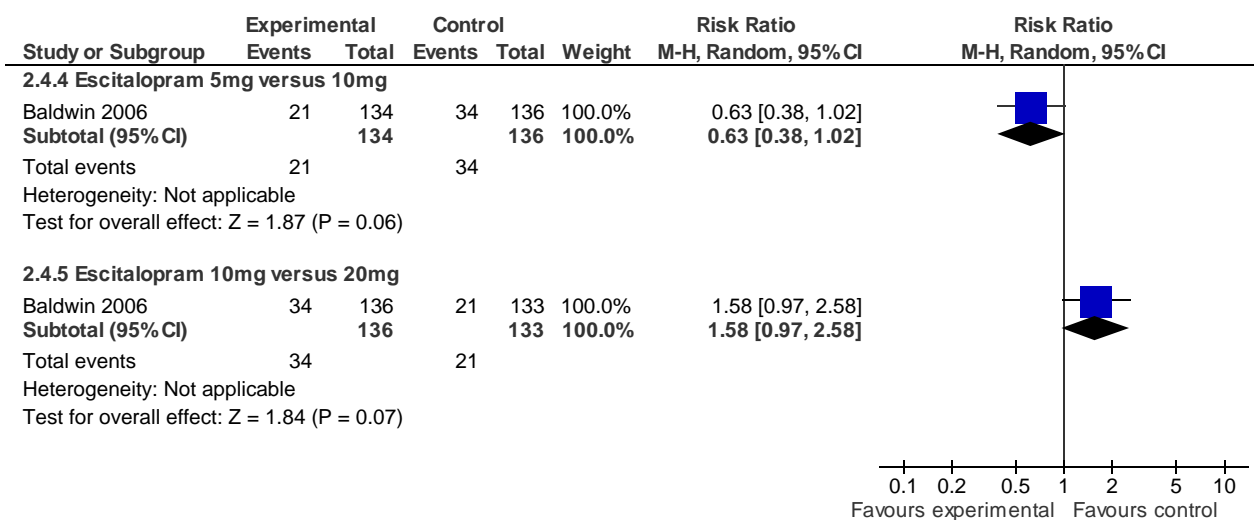


## 6 Comparing the Effectiveness of Different Dosages

### 2.3 Fatigue

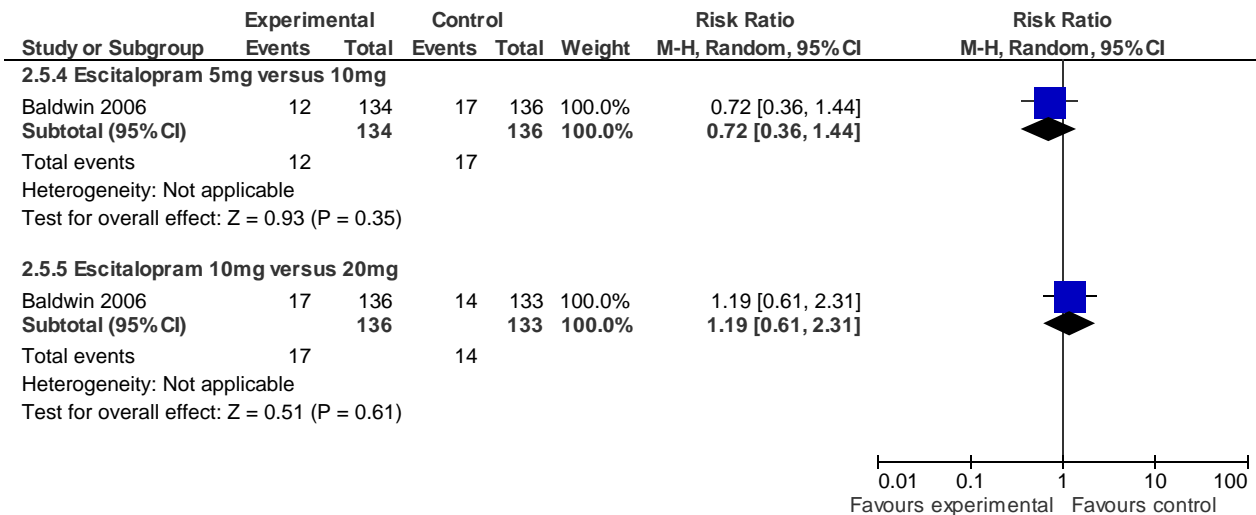


### 2.4 Headache

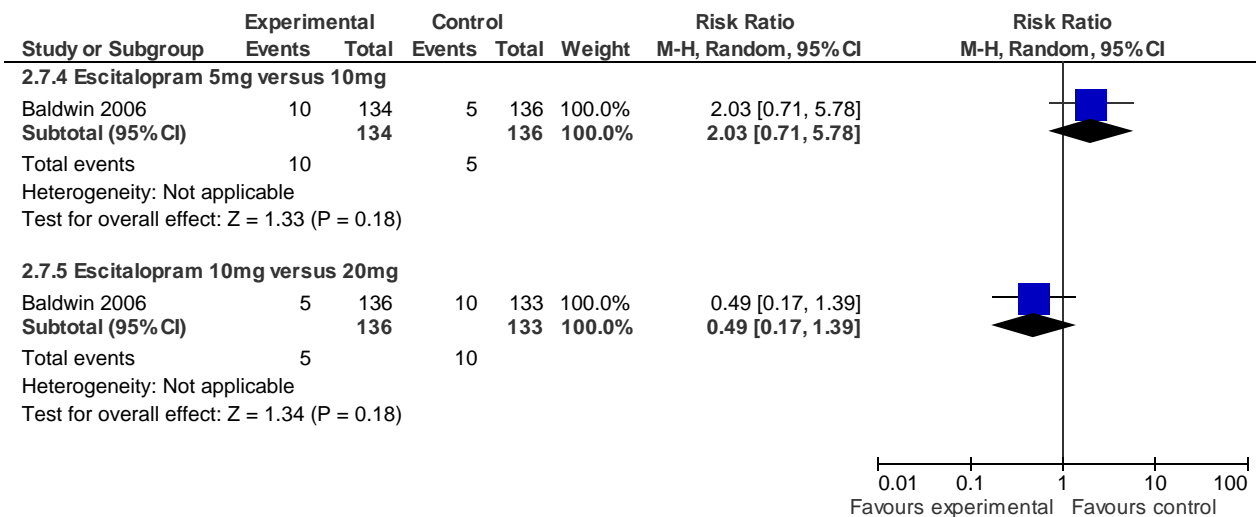


## 6 Comparing the Effectiveness of Different Dosages

### 2.5 Insomnia

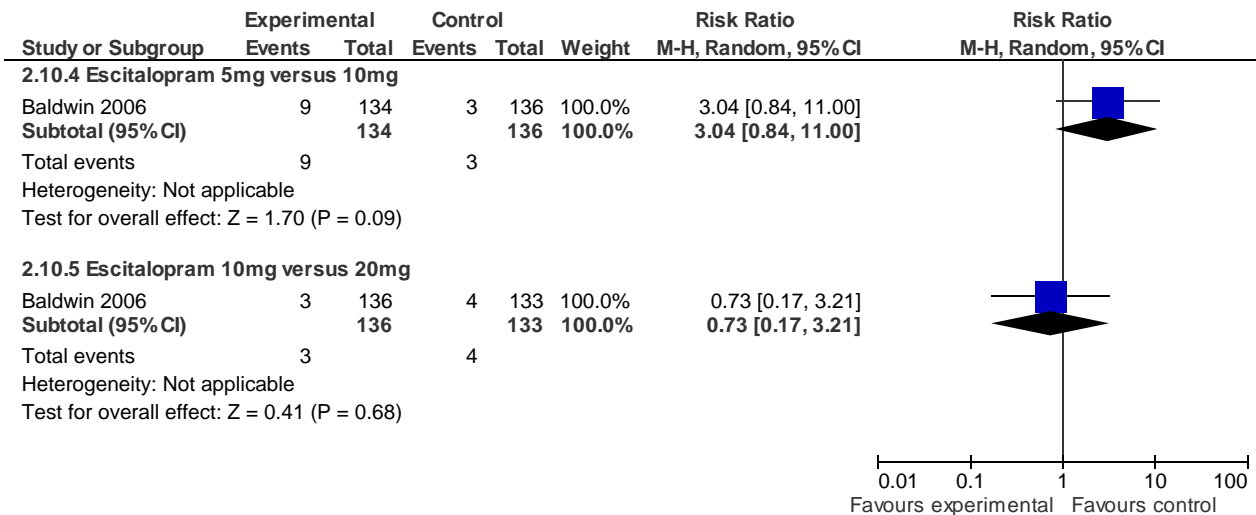


### 2.7 Somnolence

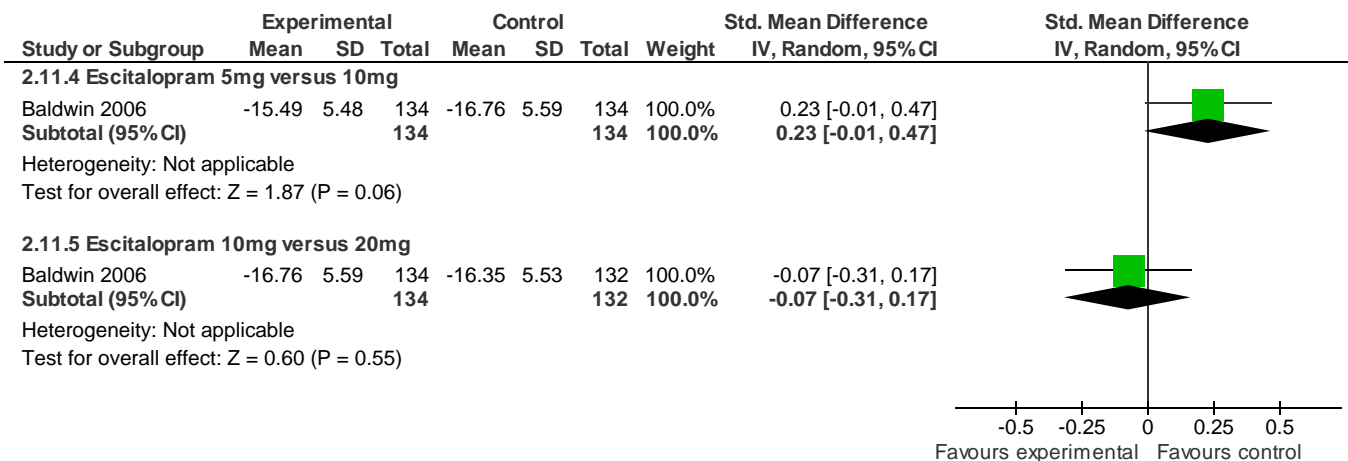


## 6 Comparing the Effectiveness of Different Dosages

### 2.10 Anxiety

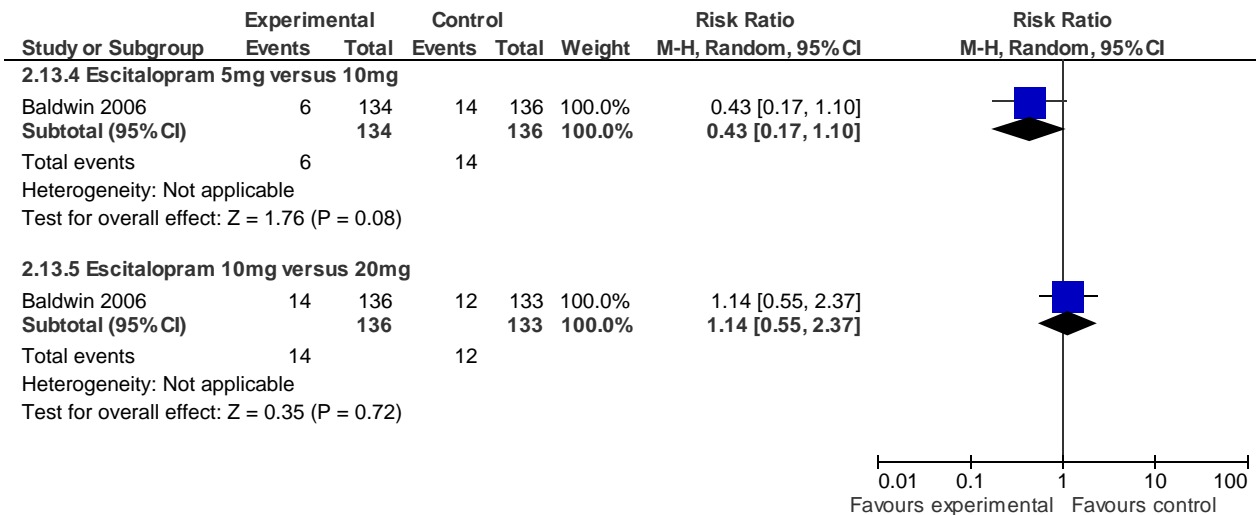


### 2.11 HAM-A



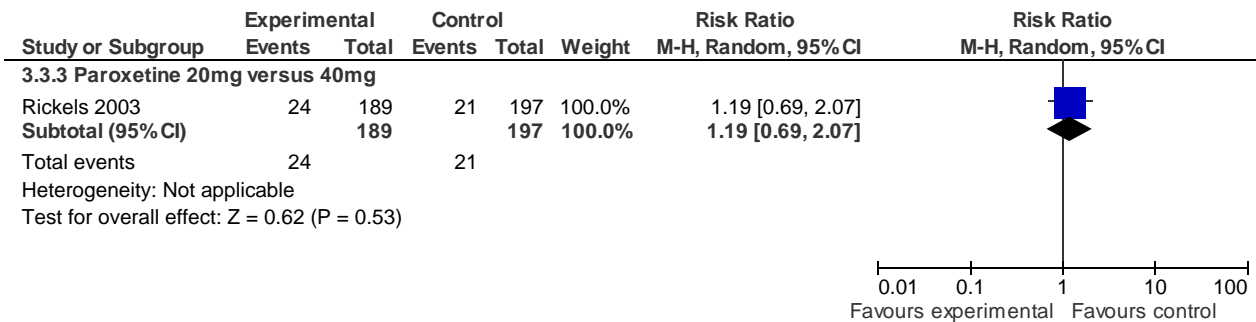
## 6 Comparing the Effectiveness of Different Dosages

### 2.13 Dizziness

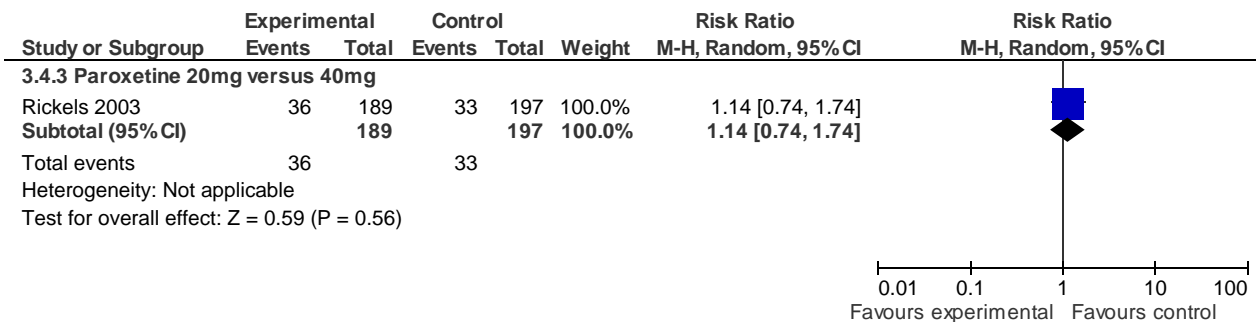


### 3 Paroxetine

#### 3.3 Decreased libido

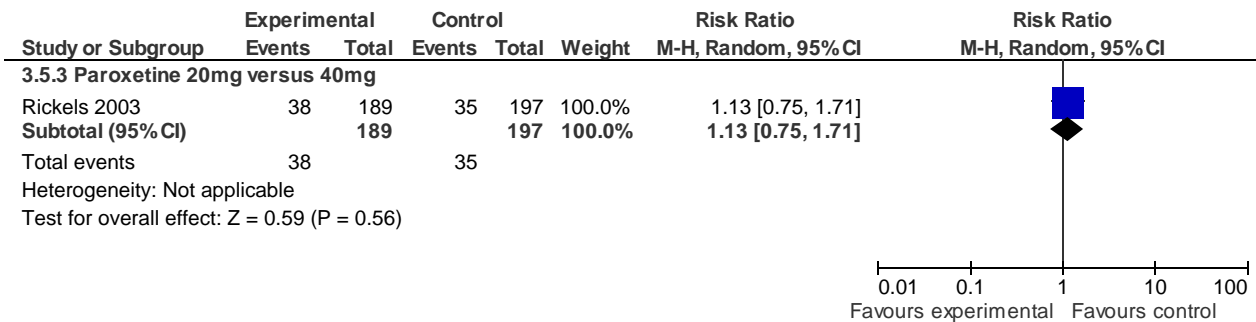


#### 3.4 Nausea

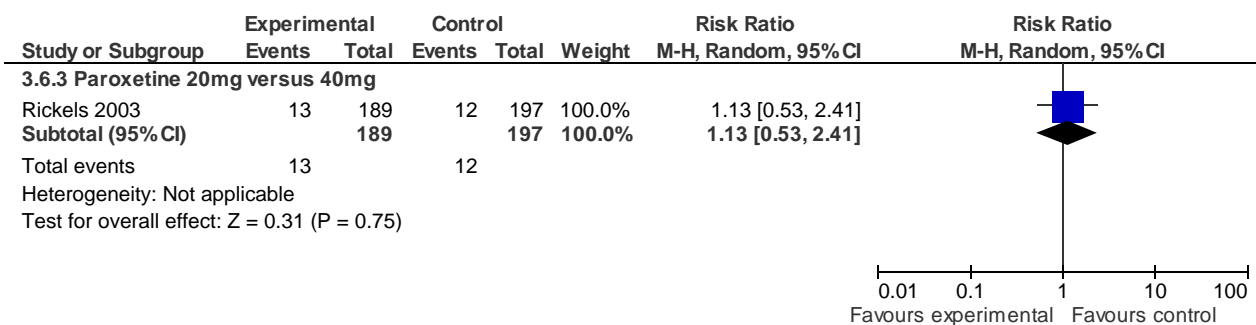


## 6 Comparing the Effectiveness of Different Dosages

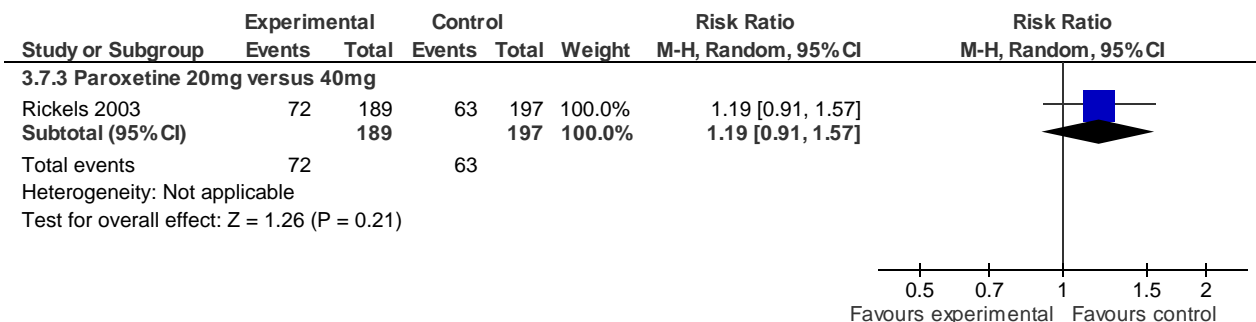
### 3.5 Somnolence



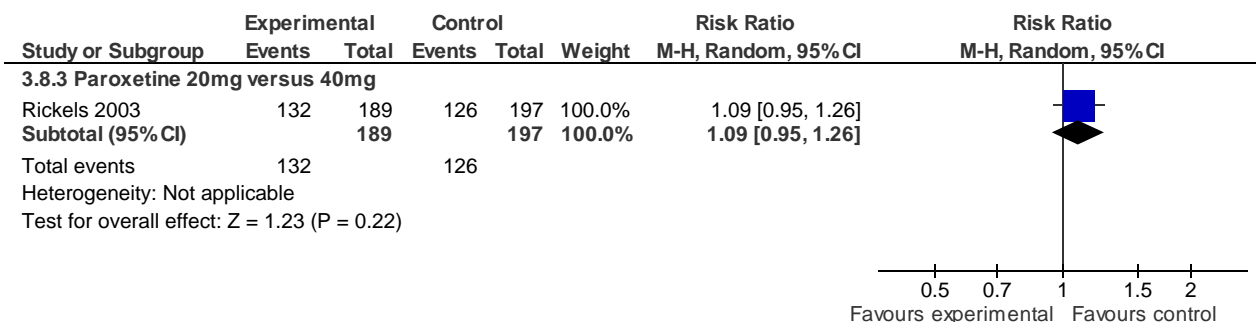
### 3.6 Decreased appetite



### 3.7 Non-response

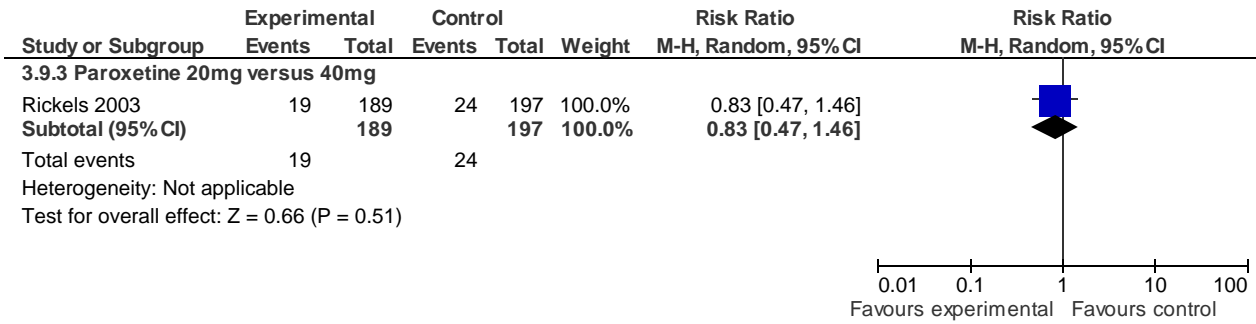


### 3.8 Non-remission

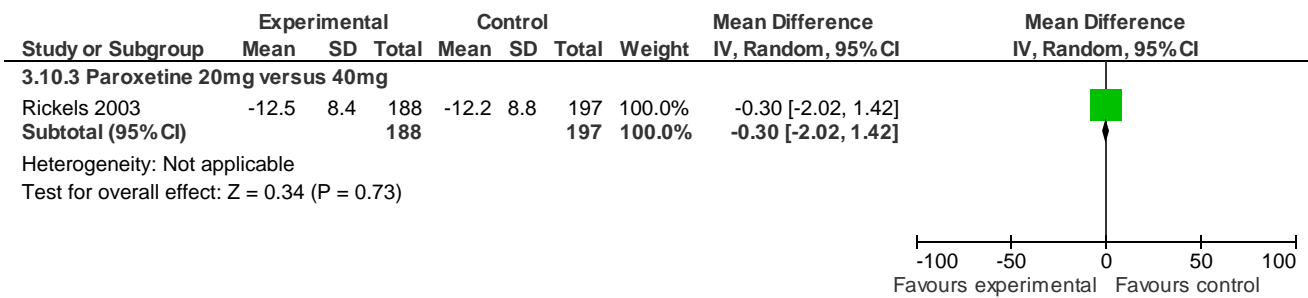


## 6 Comparing the Effectiveness of Different Dosages

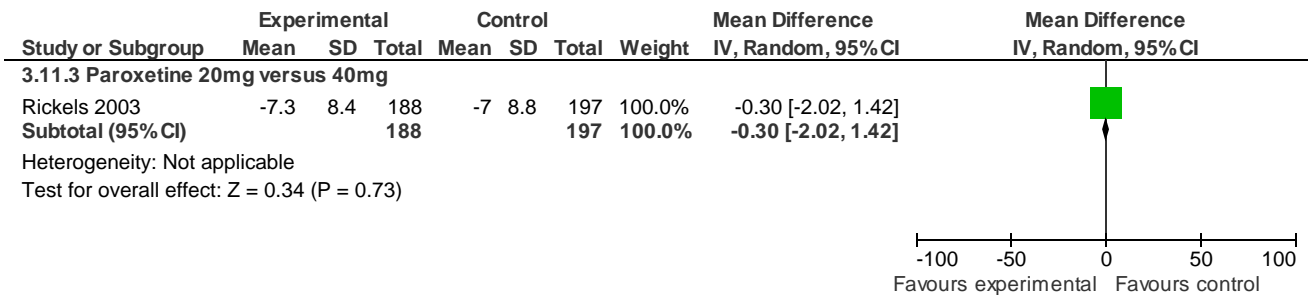
### 3.9 Discontinuation due to Adverse Events



### 3.10 HAM-A



### 3.11 HADS-A

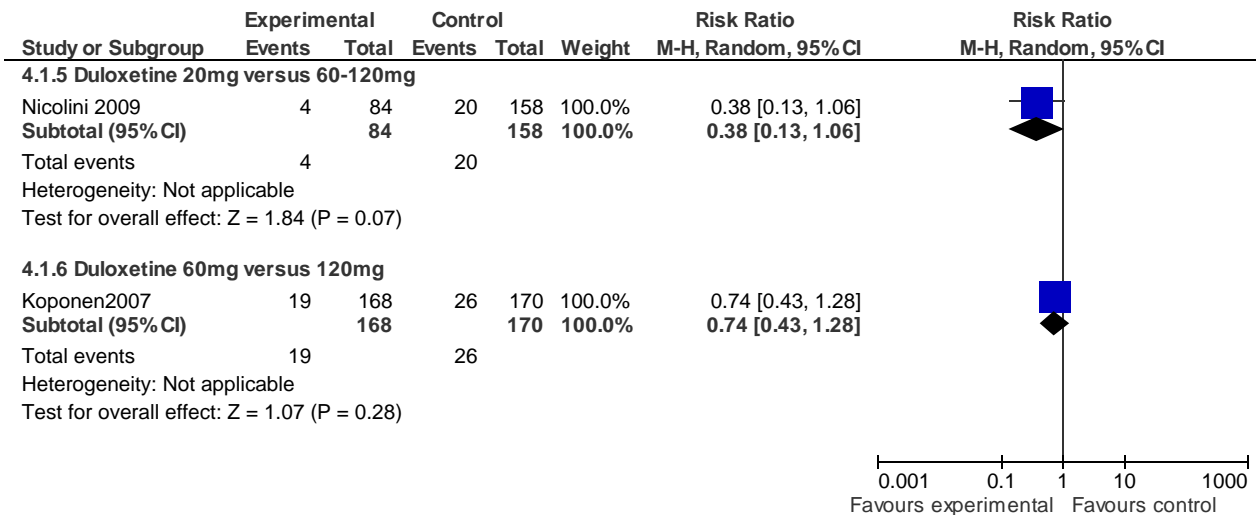


## 4 Duloxetine

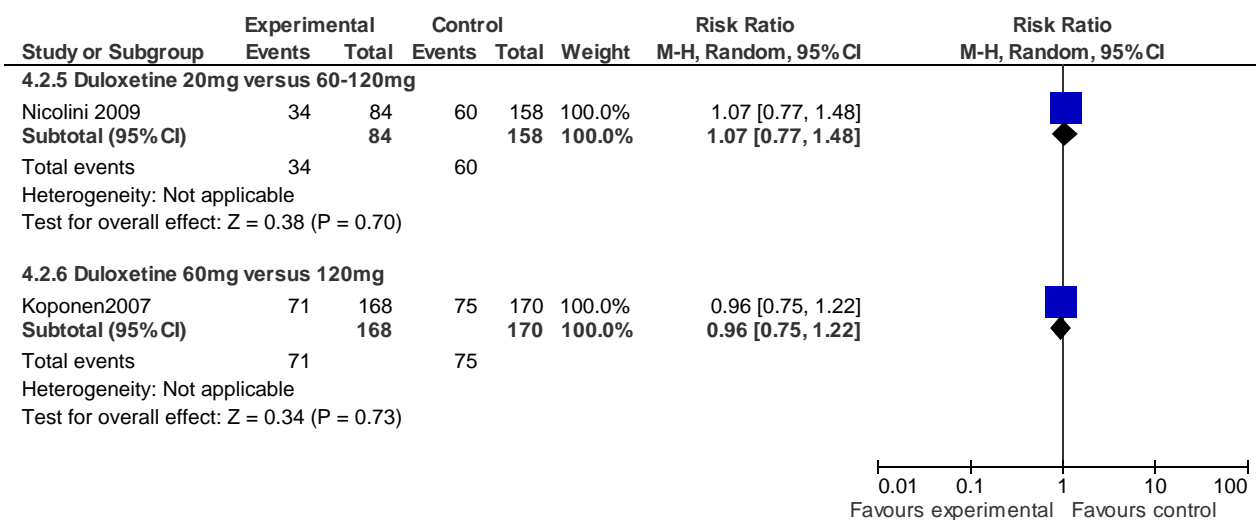


## 6 Comparing the Effectiveness of Different Dosages

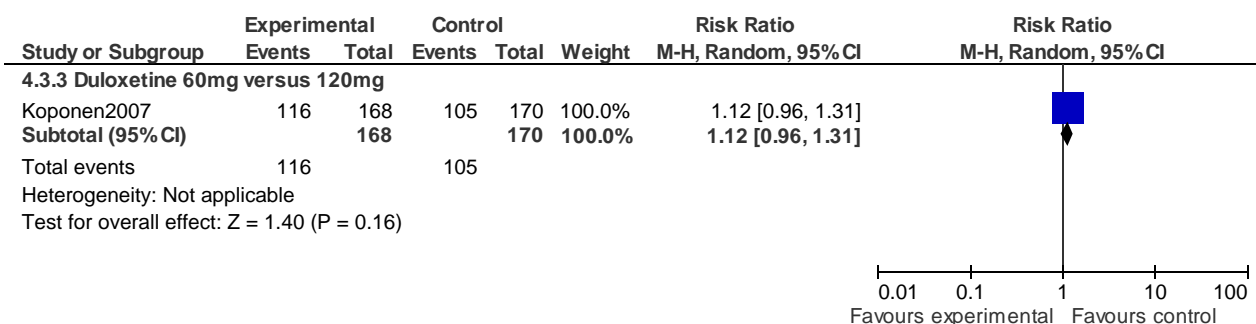
### 4.1 Discontinuation due to Adverse Events



### 4.2 Non-response

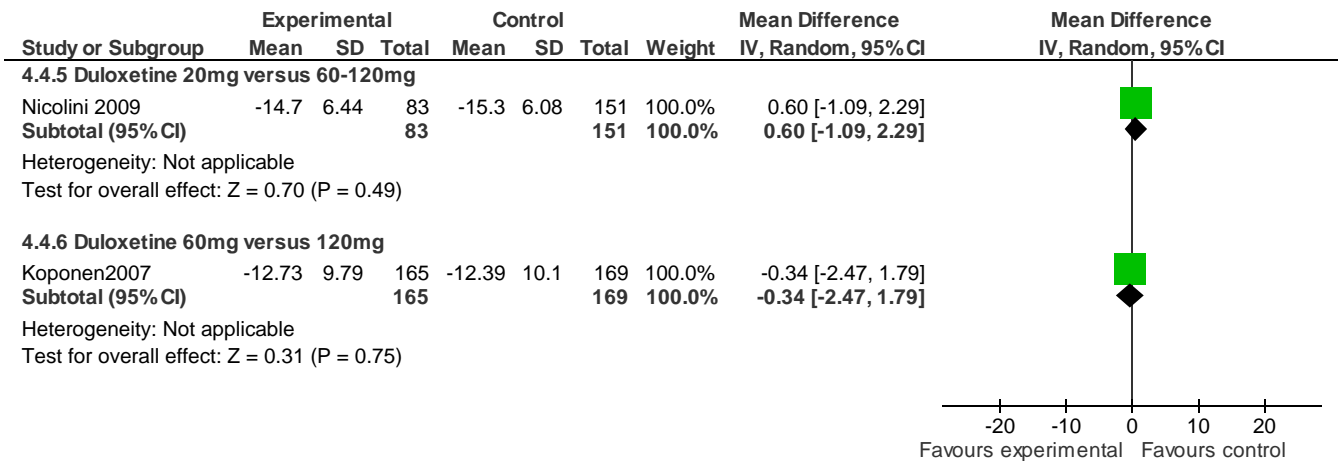


### 4.3 Non-remission

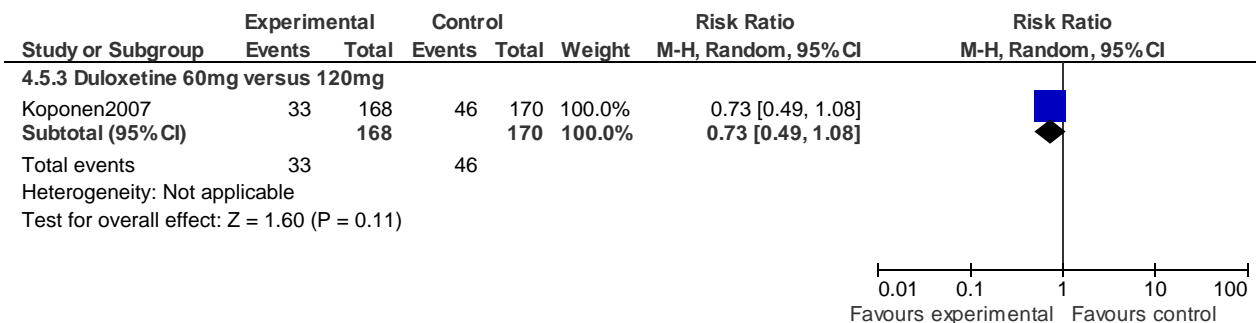


## 6 Comparing the Effectiveness of Different Dosages

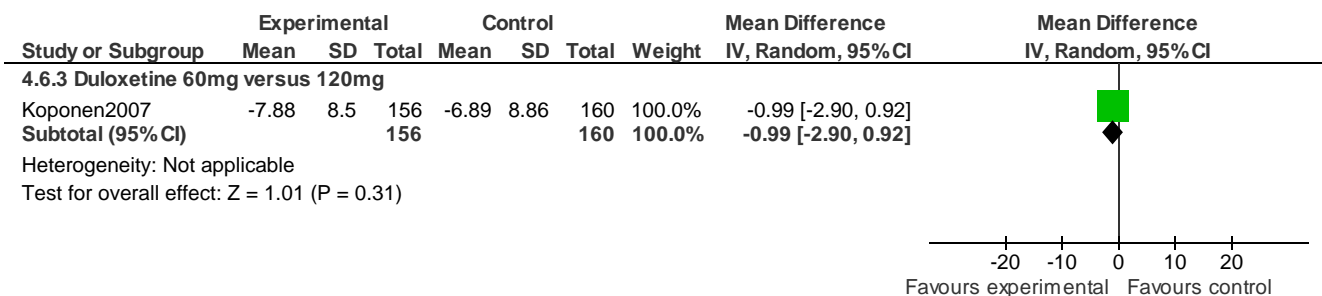
### 4.4 HAM-A



### 4.5 Discontinuation due to Any Reason

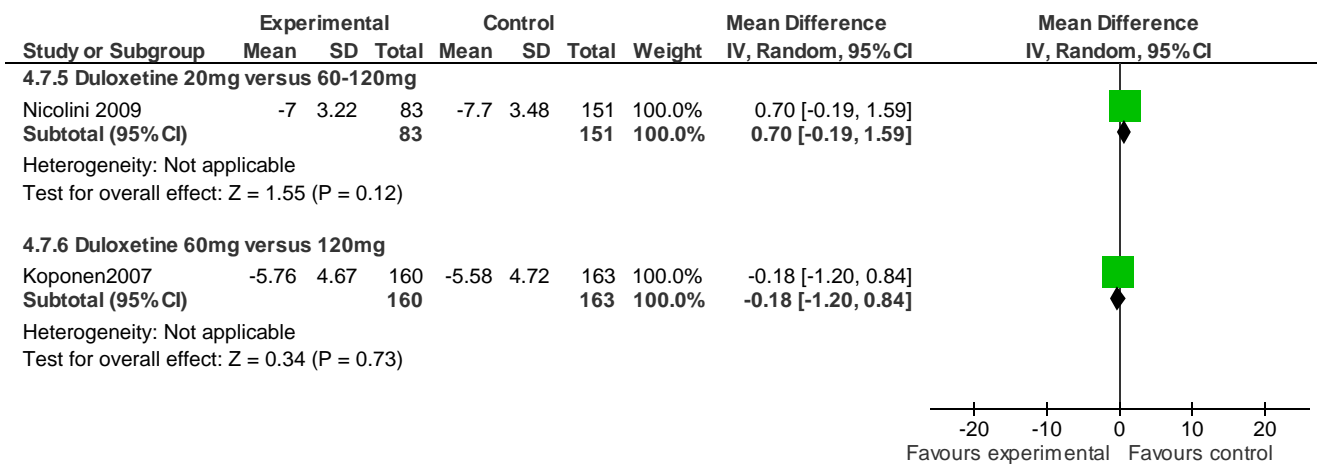


### 4.6 Sheehan Disability Scale

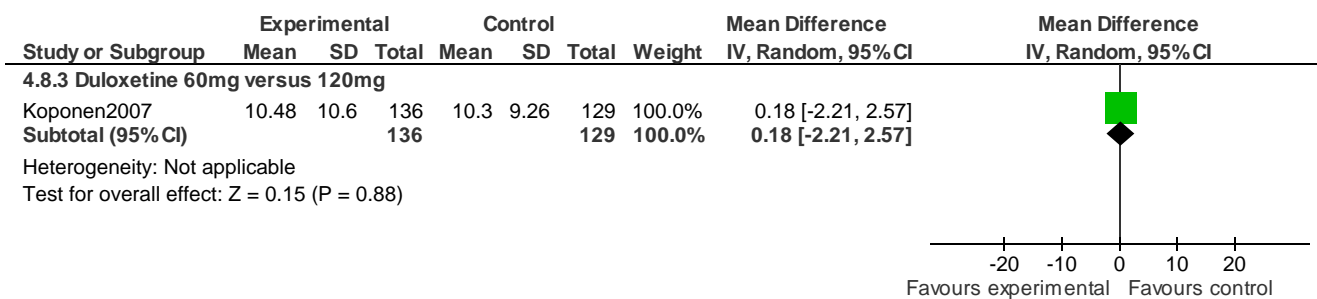


## 6 Comparing the Effectiveness of Different Dosages

### 4.7 HADS-A



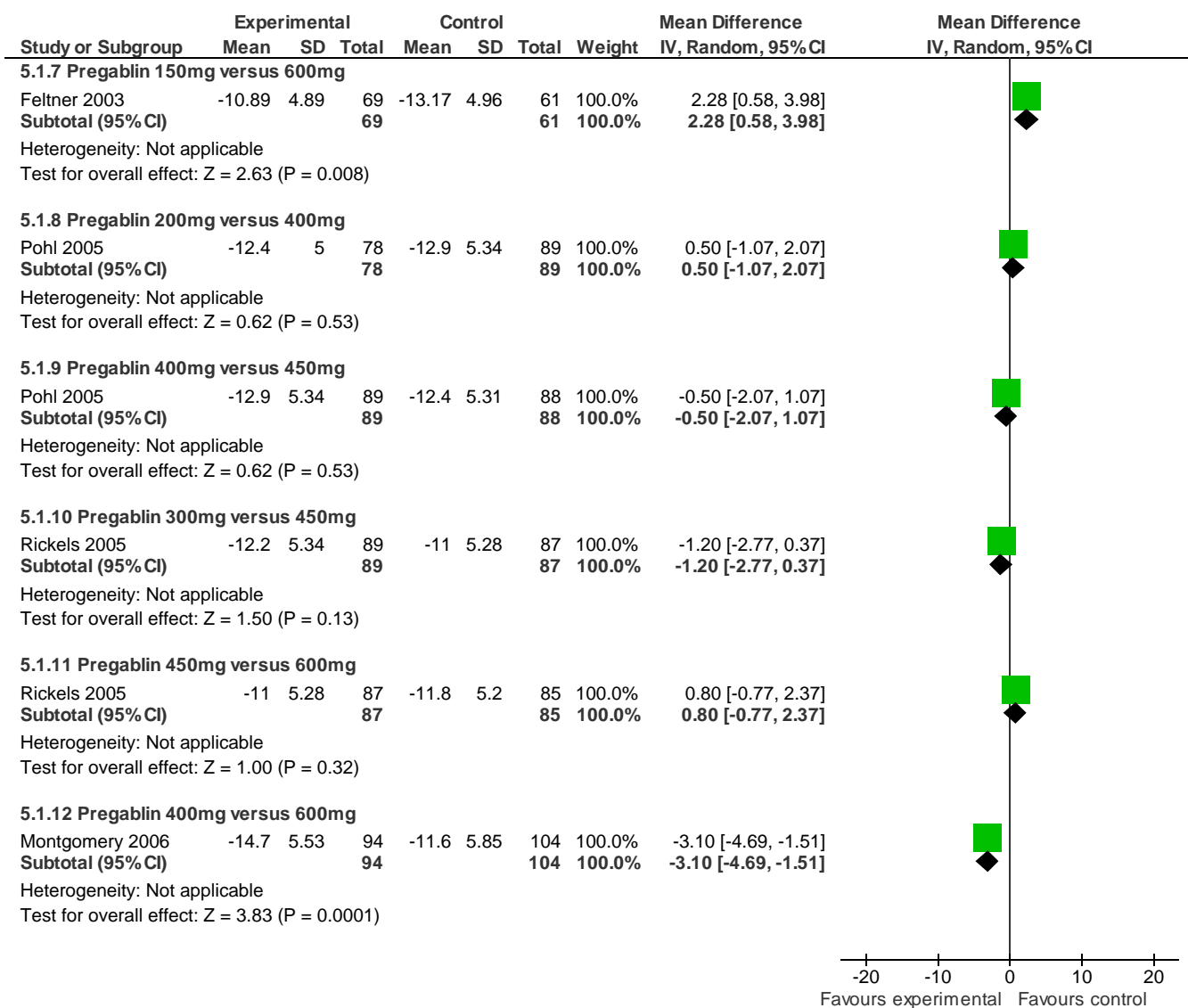
### 4.8 Q-LES-Q-SF



### 5 Pregablin

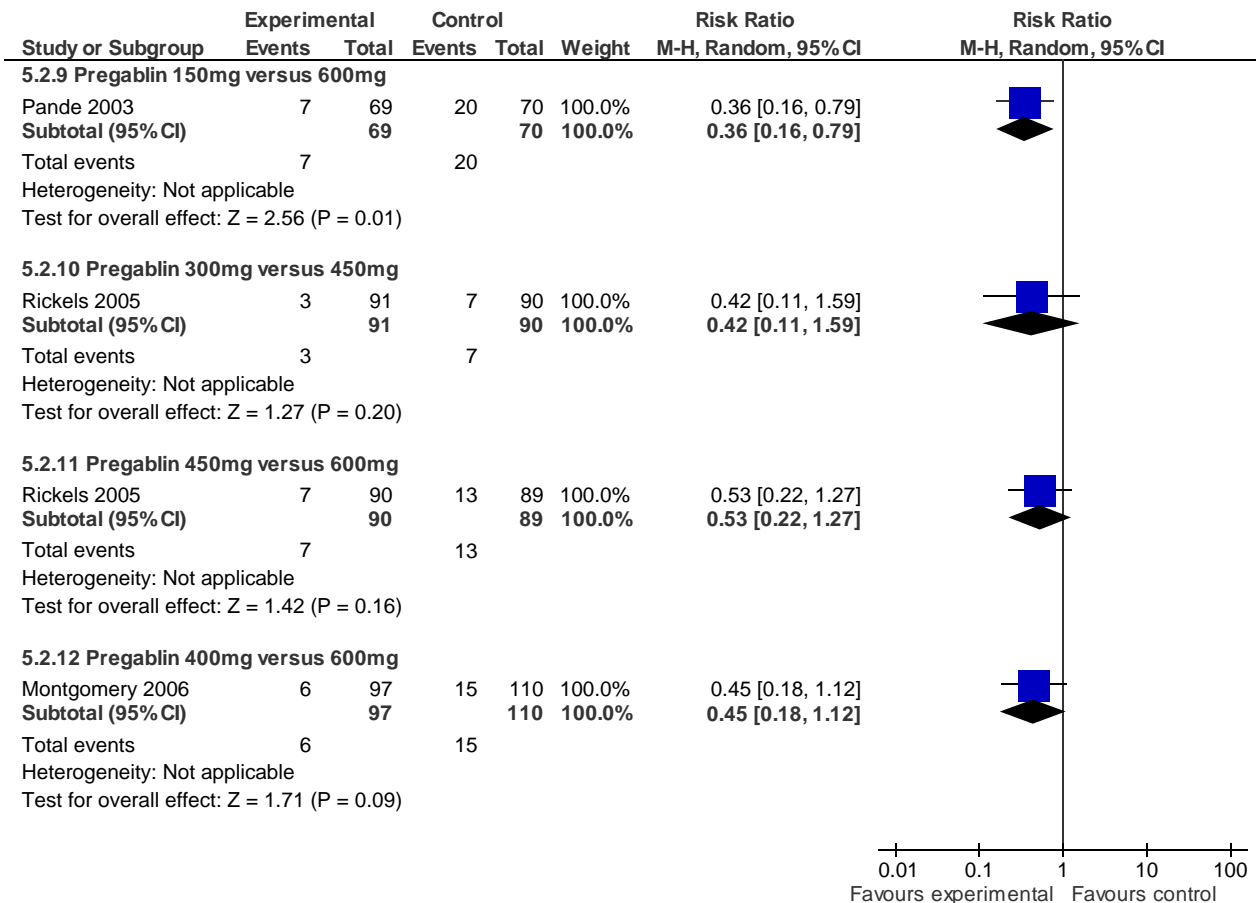
## 6 Comparing the Effectiveness of Different Dosages

### 5.1 HAM-A

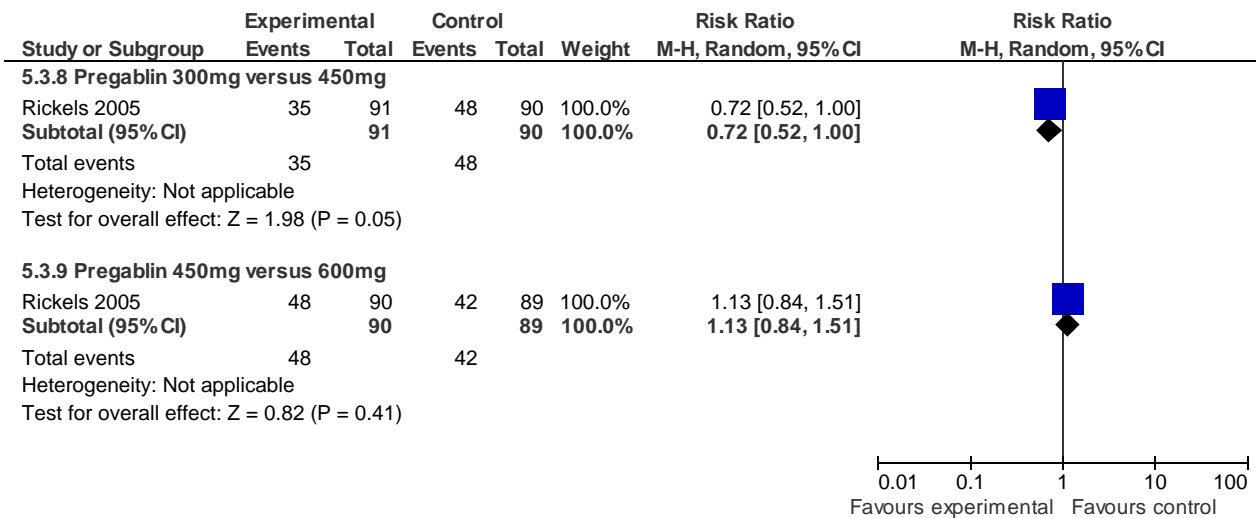


## 6 Comparing the Effectiveness of Different Dosages

### 5.2 Discontinuation due to Adverse Events

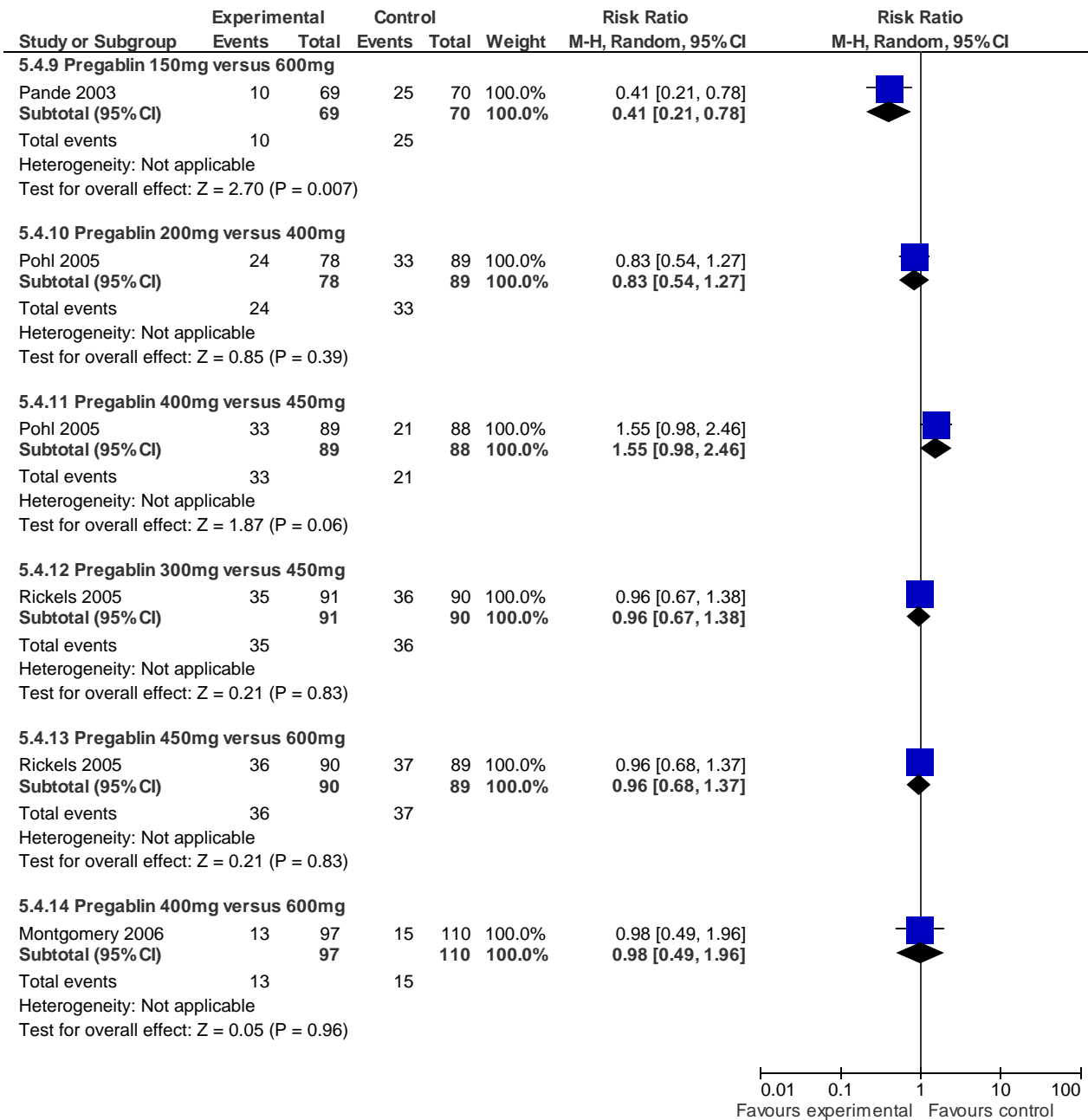


### 5.3 Non Response



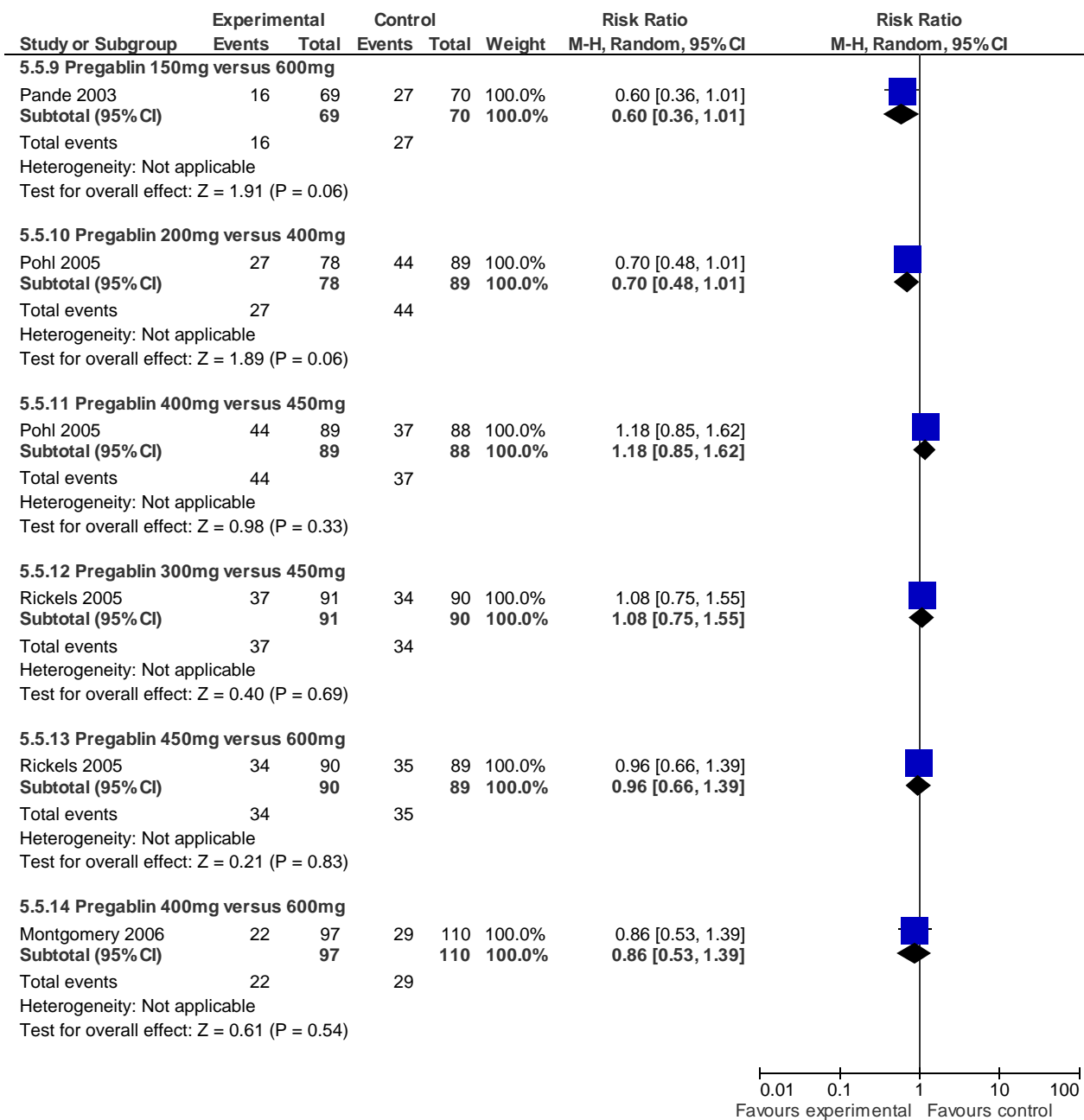
## 6 Comparing the Effectiveness of Different Dosages

### 5.4 Somnolence



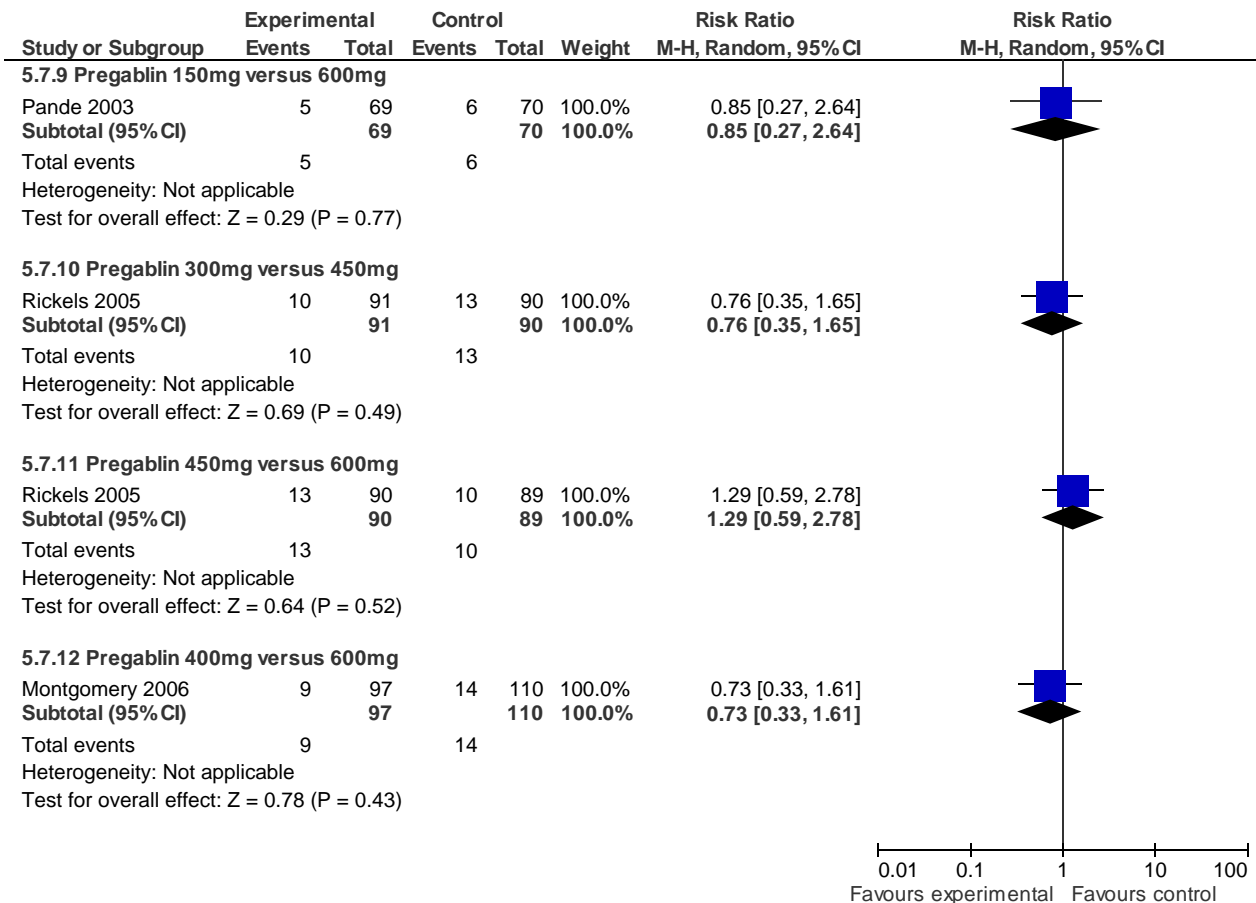
## 6 Comparing the Effectiveness of Different Dosages

### 5.5 Dizziness

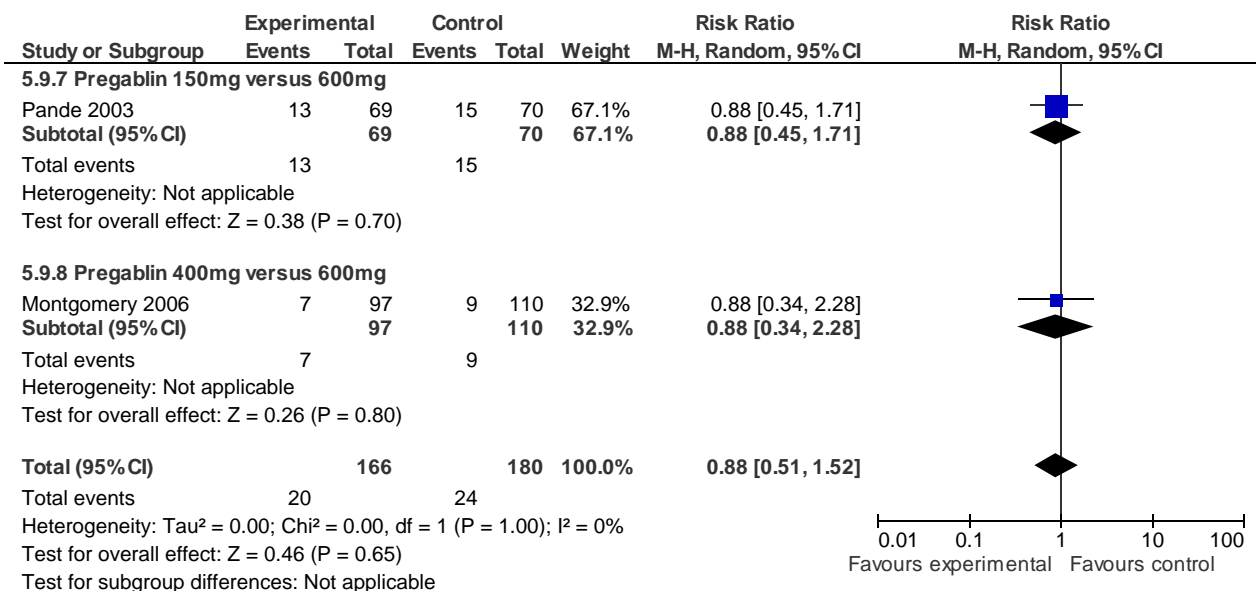


## 6 Comparing the Effectiveness of Different Dosages

### 5.7 Nausea



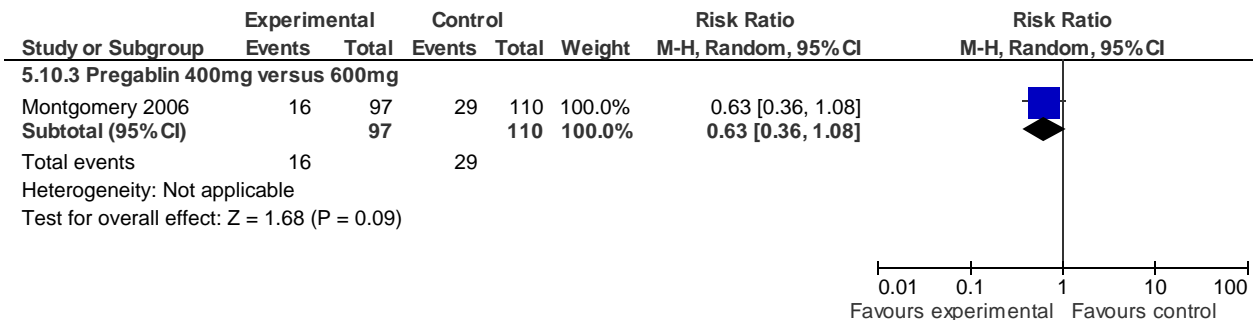
### 5.9 Headache



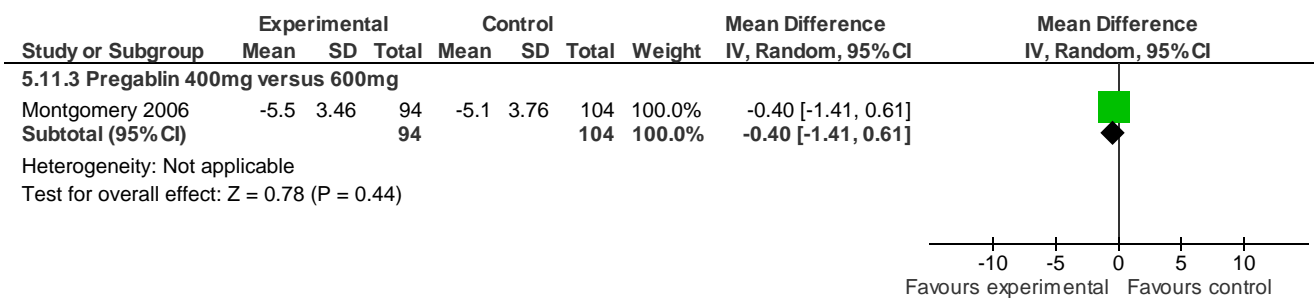


## 6 Comparing the Effectiveness of Different Dosages

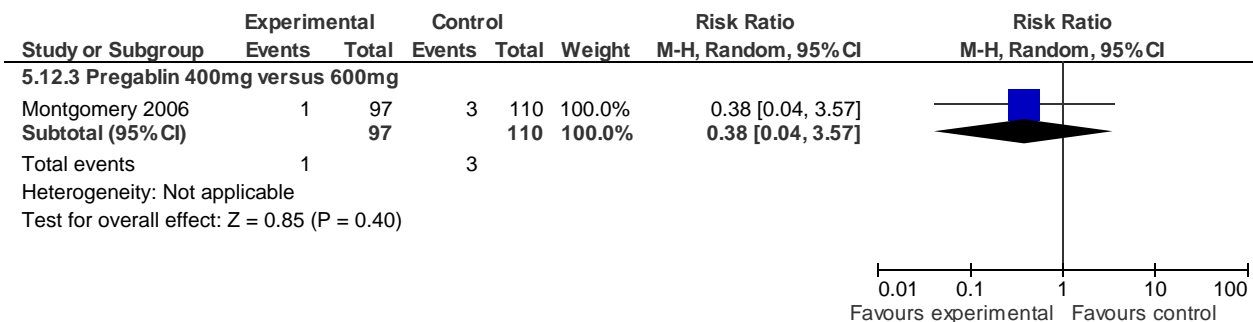
### 5.10 Discontinuation for any reason



### 5.11 HADS-A



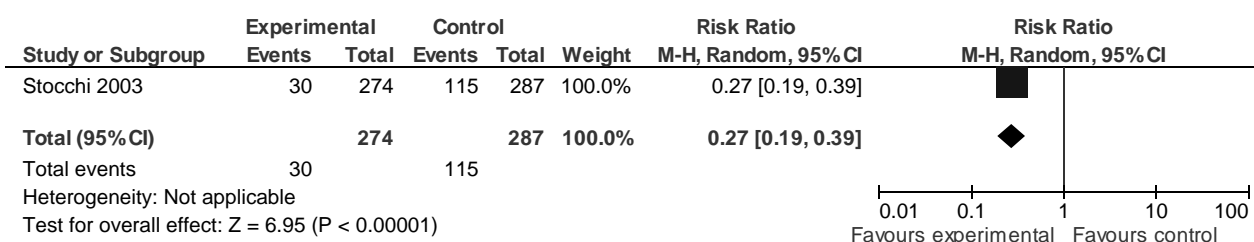
### 5.12 Insomnia



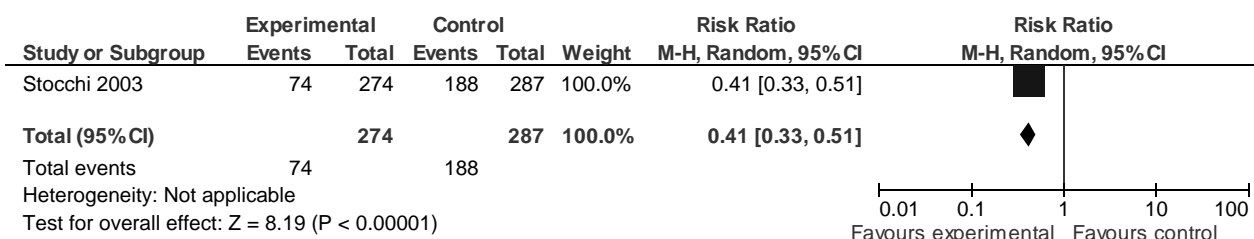
## 7 Maintenance Treatment

### 1 Paroxetine versus Placebo

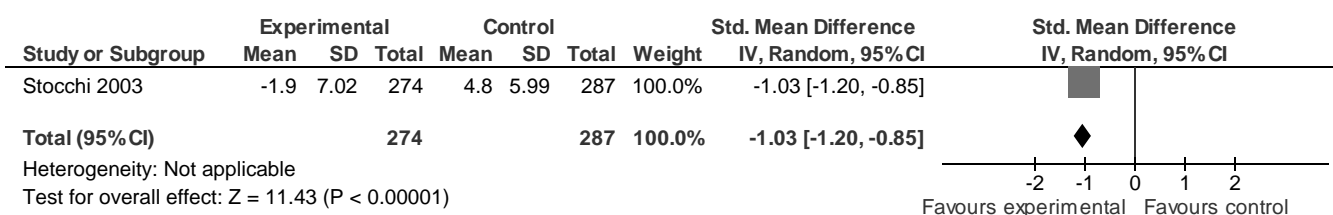
#### 1.1 Relapse



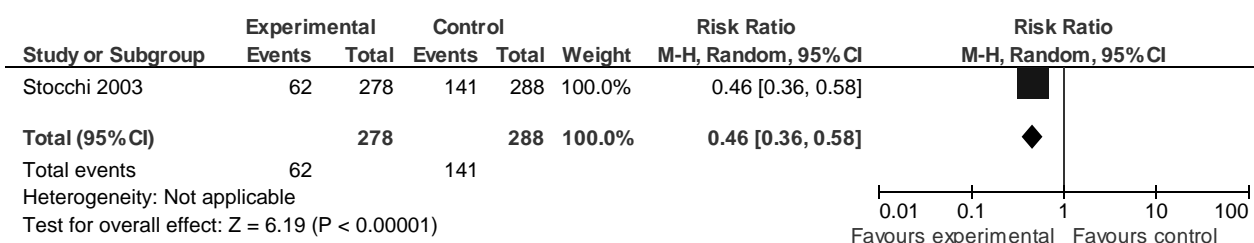
#### 1.2 Non-remission



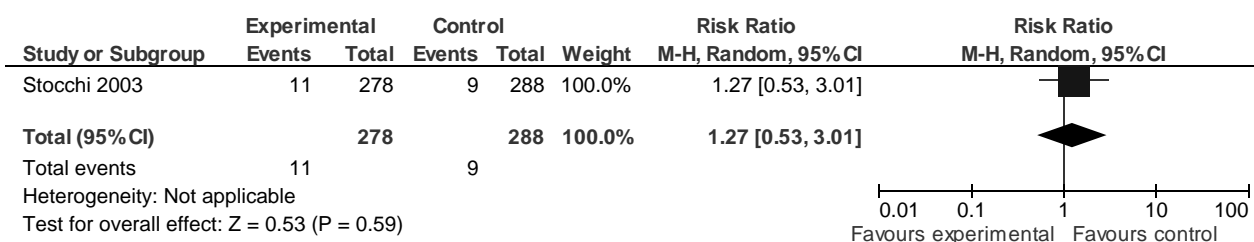
#### 1.3 HAM-A



#### 1.4 Discontinuation for any reason



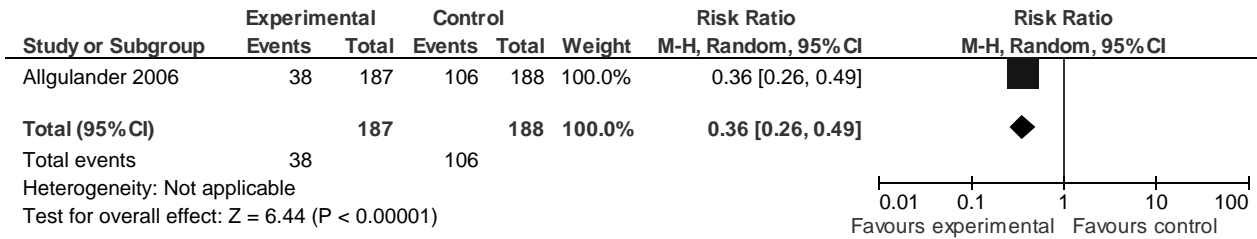
#### 1.5 Discontinuation due to adverse events



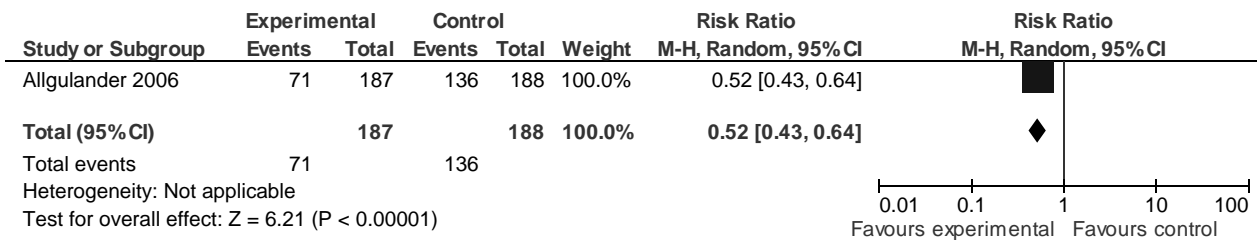
### 2 Escitalopram versus Placebo

## 7 Maintenance Treatment

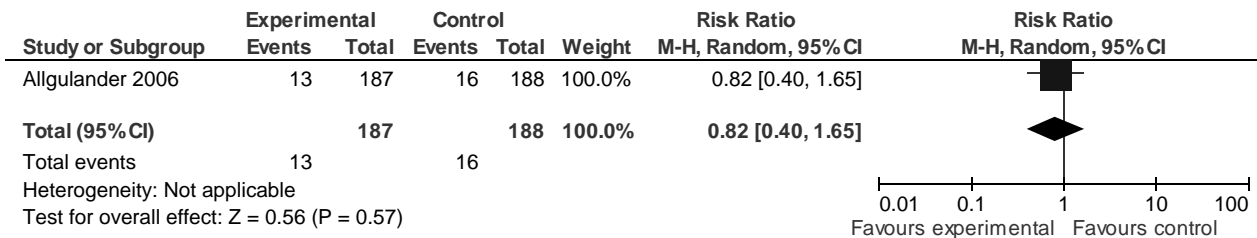
### 2.1 Relapse



### 2.2 Discontinuation for any reason

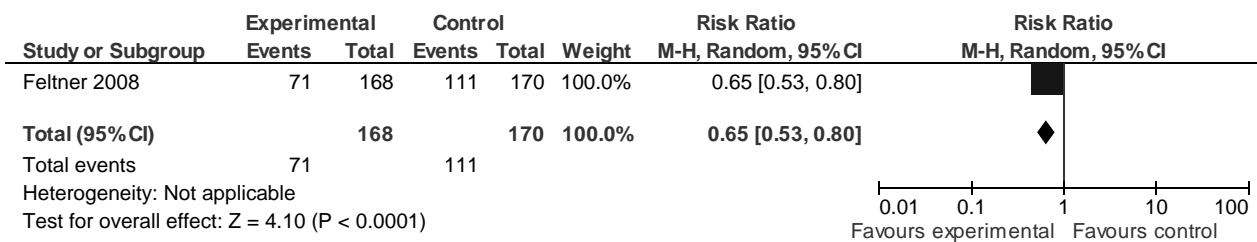


### 2.3 Discontinuation due to adverse events

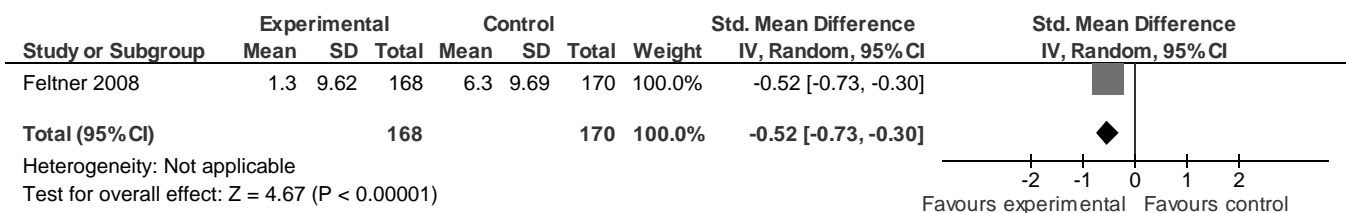


## 3 Pregabalin versus Placebo

### 3.1 Relapse

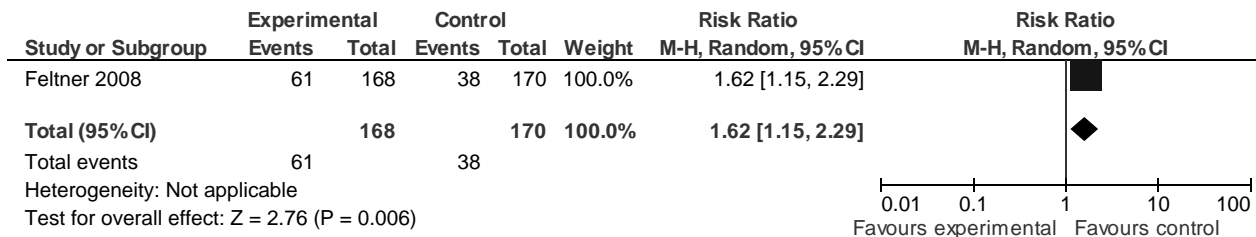


### 3.2 HAM-A

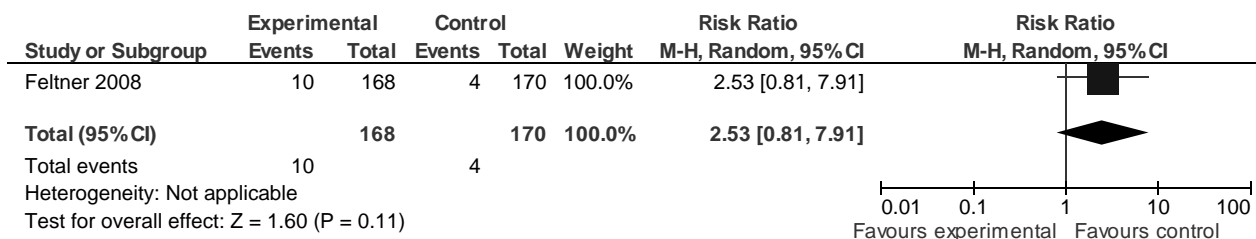


## 7 Maintenance Treatment

### 3.3 Discontinuation for any reason

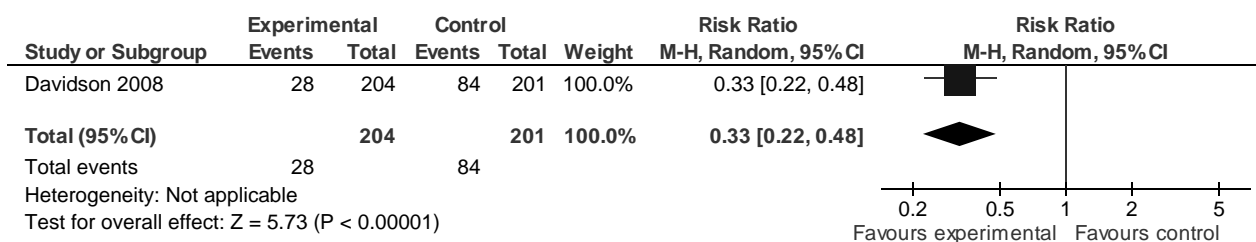


### 3.4 Discontinuation due to adverse events

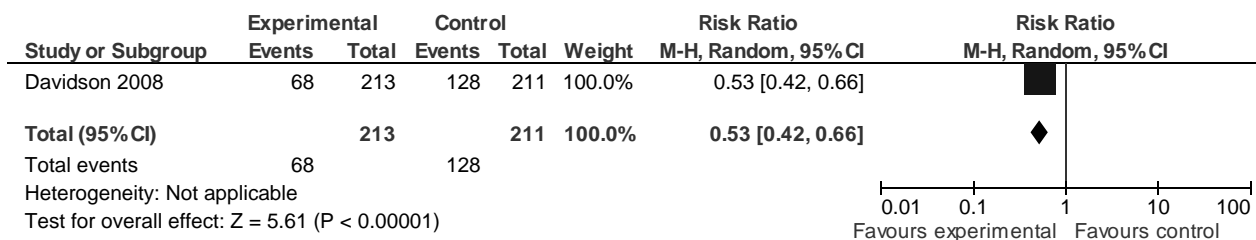


## 4 Duloxetine versus Placebo

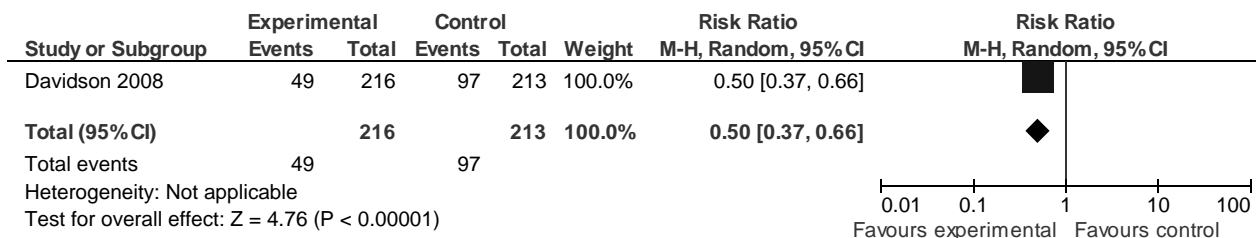
### 4.1 Relapse



### 4.2 Non-remission

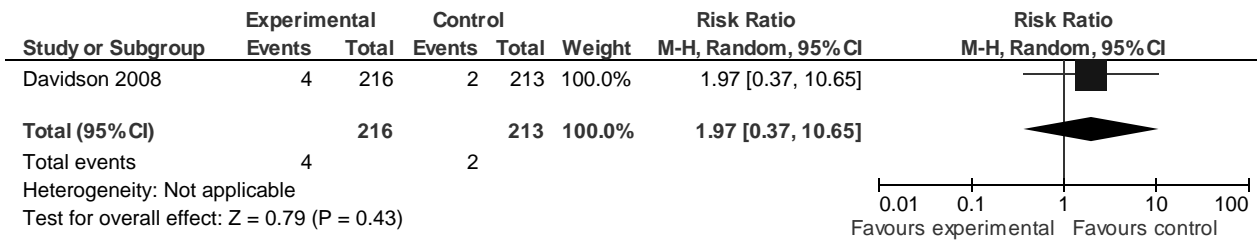


### 4.3 Discontinuation for any reason

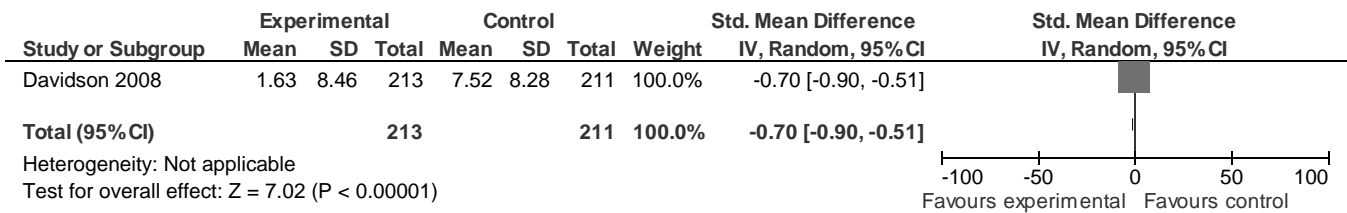


## 7 Maintenance Treatment

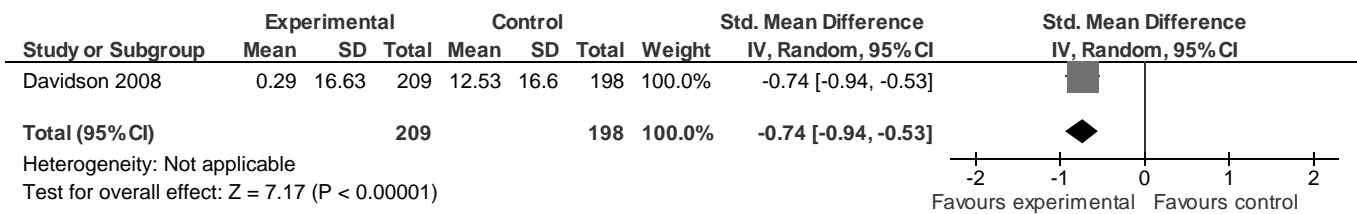
### 4.4 Discontinuation due to adverse events



### 4.5 HAM-A



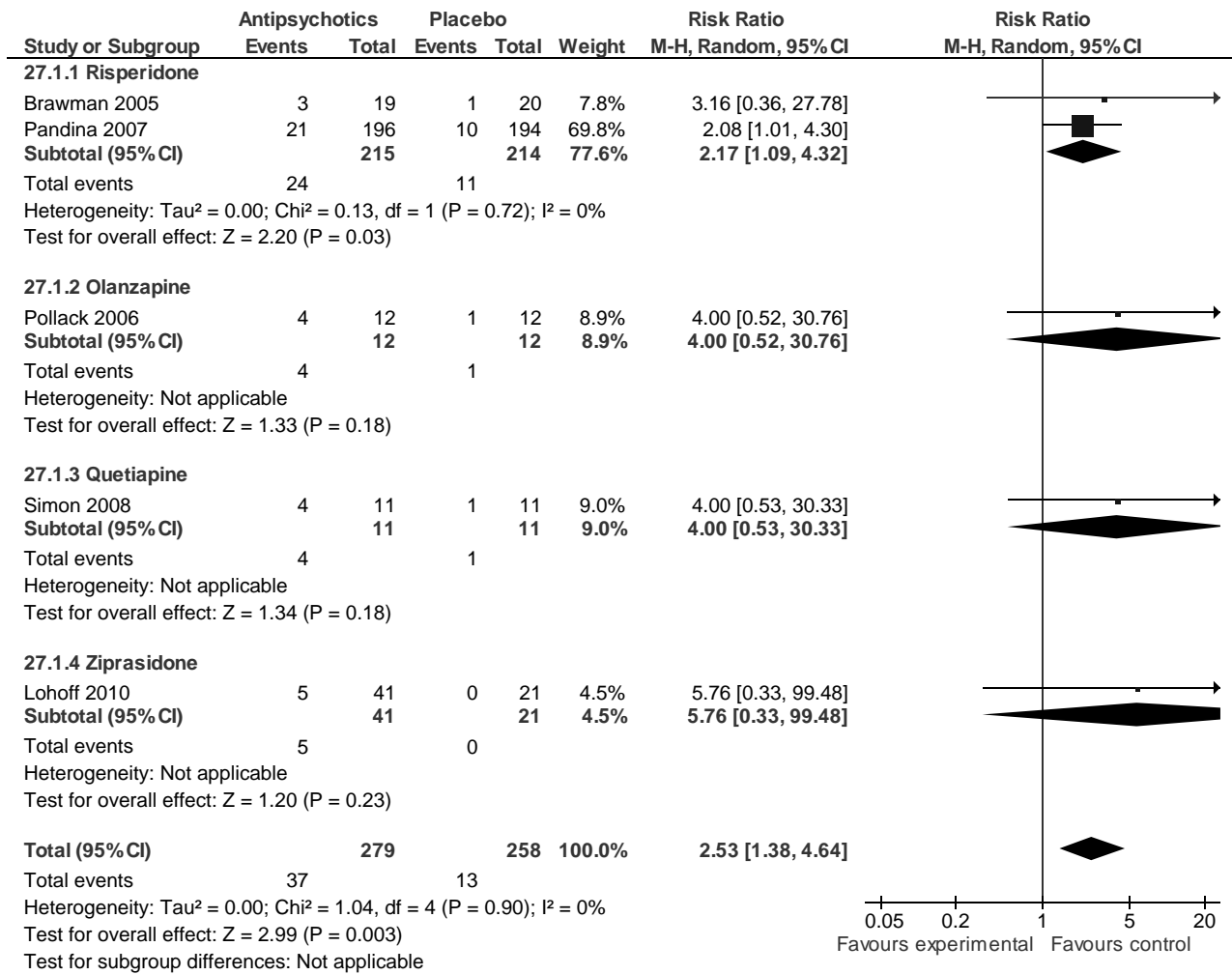
### 4.6 Q-LES-Q-SF



# 8 Augmentation

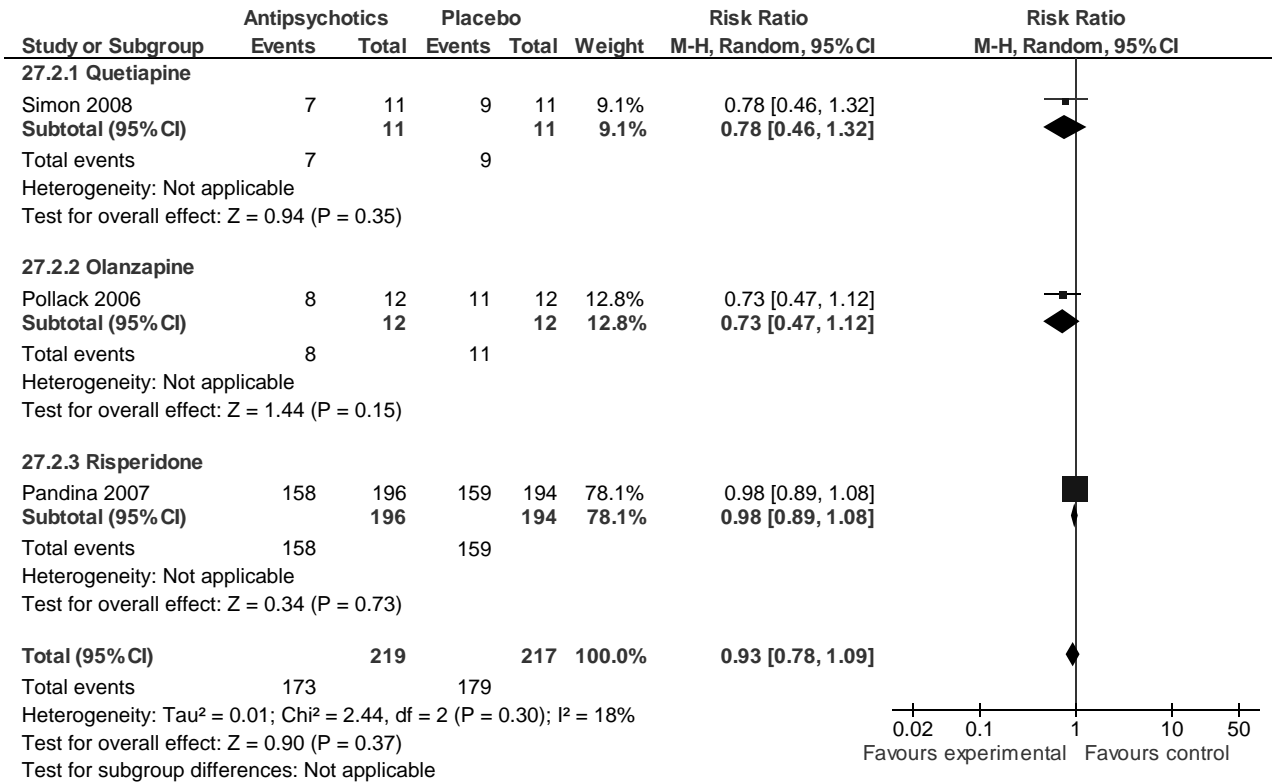
## 27 Augmentation: Antipsychotics versus Placebo

### 27.1 Discontinuation due to adverse events

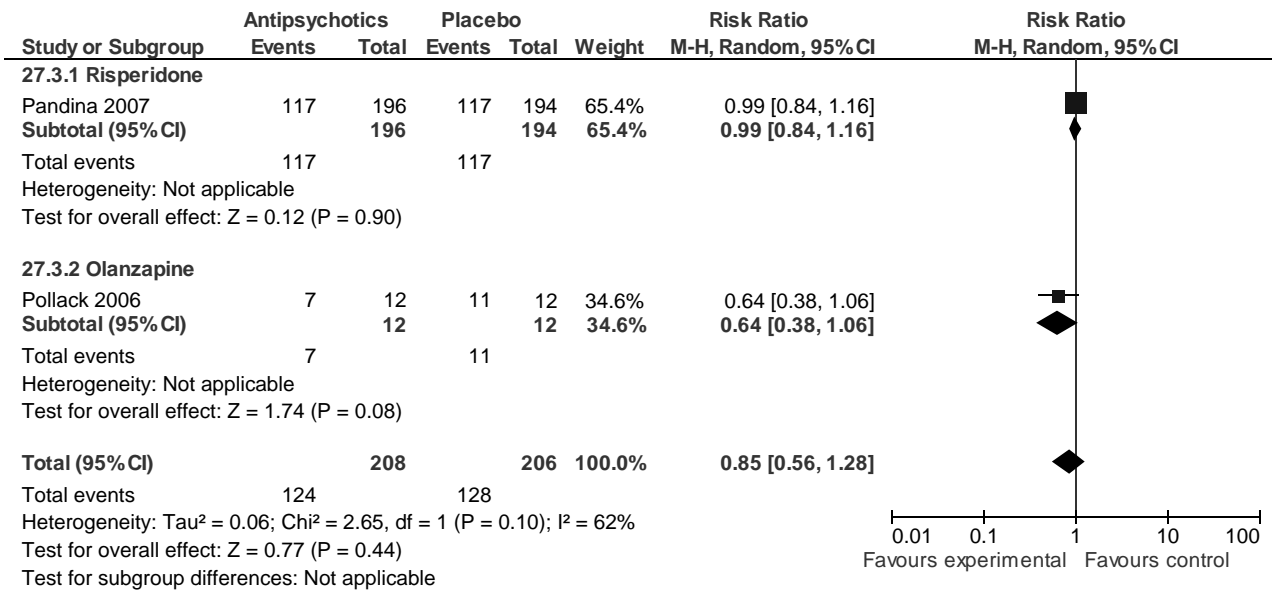


## 8 Augmentation

### 27.2 Non-remission

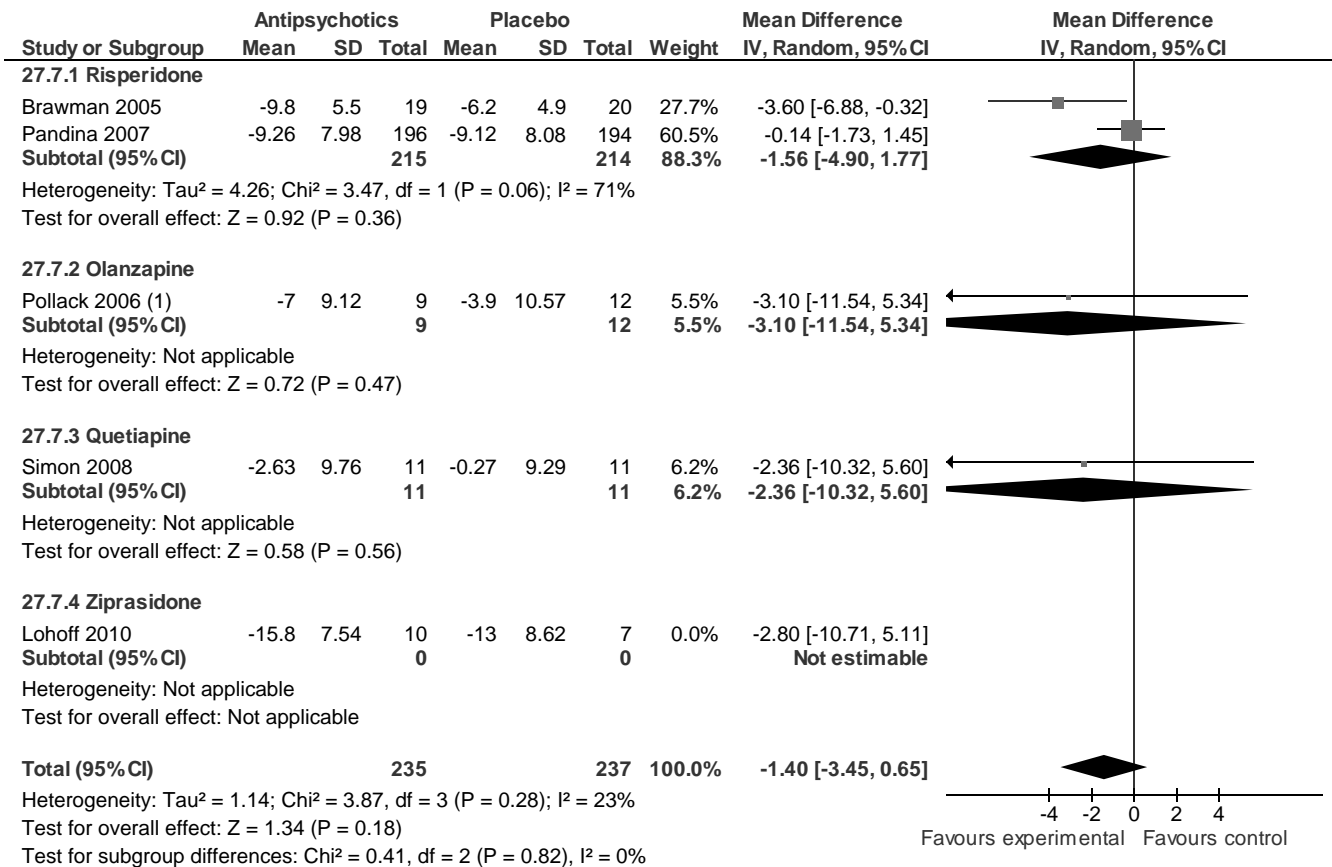


### 27.3 Non-response



# 8 Augmentation

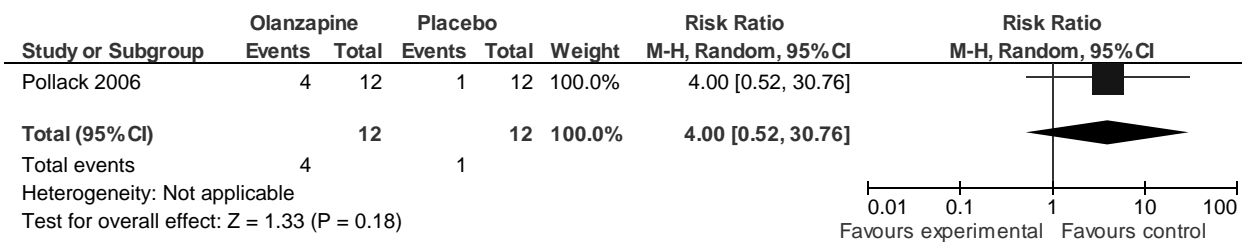
## 27.7 HAM-A



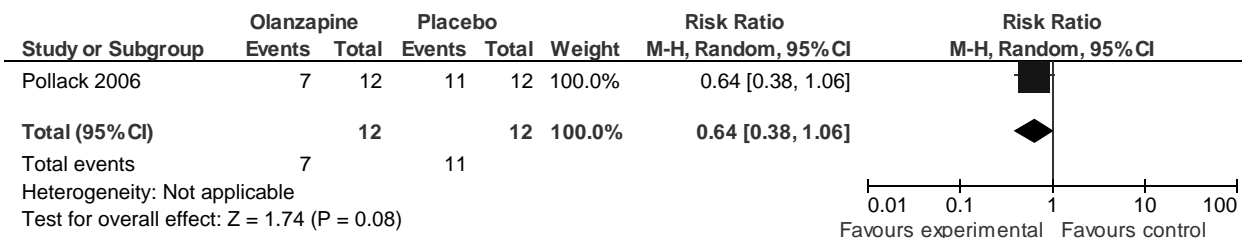
(1) change score

## 29 Augmentation: Olanzapine versus Placebo

### 29.2 Discontinuation due to adverse events



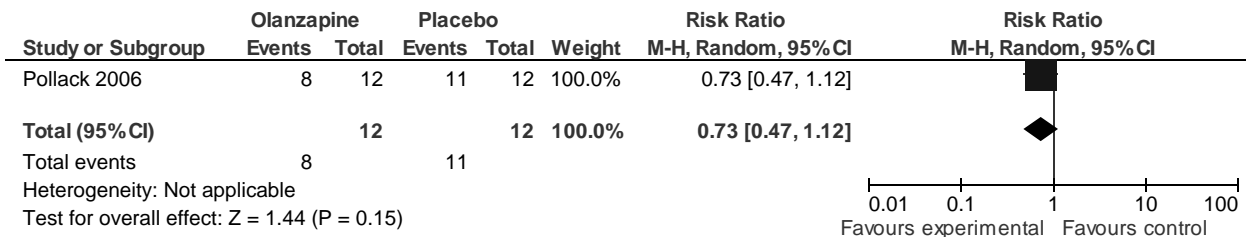
### 29.3 Non-response



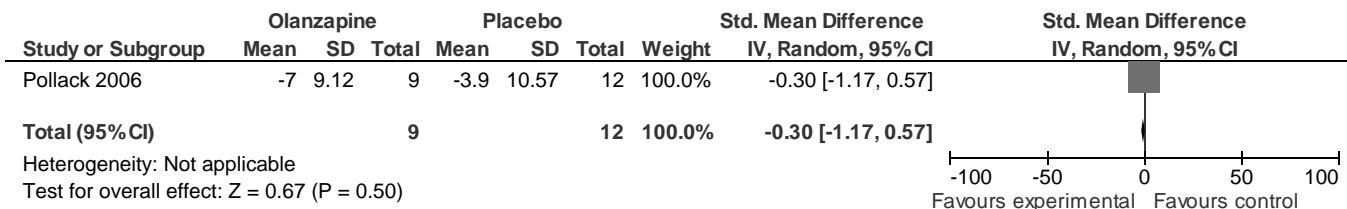


## 8 Augmentation

### 29.4 Non-remission

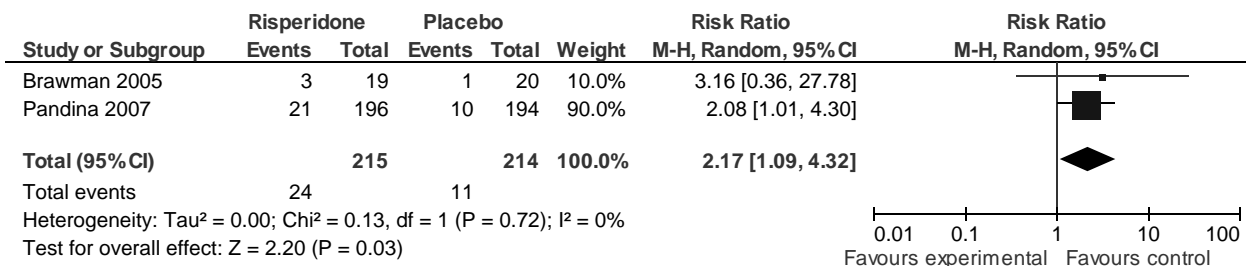


### 29.5 HAM-A



## 30 Augmentation: Risperidone versus Placebo

### 30.2 Discontinuation due to adverse events



### 30.6 HAM-A

