

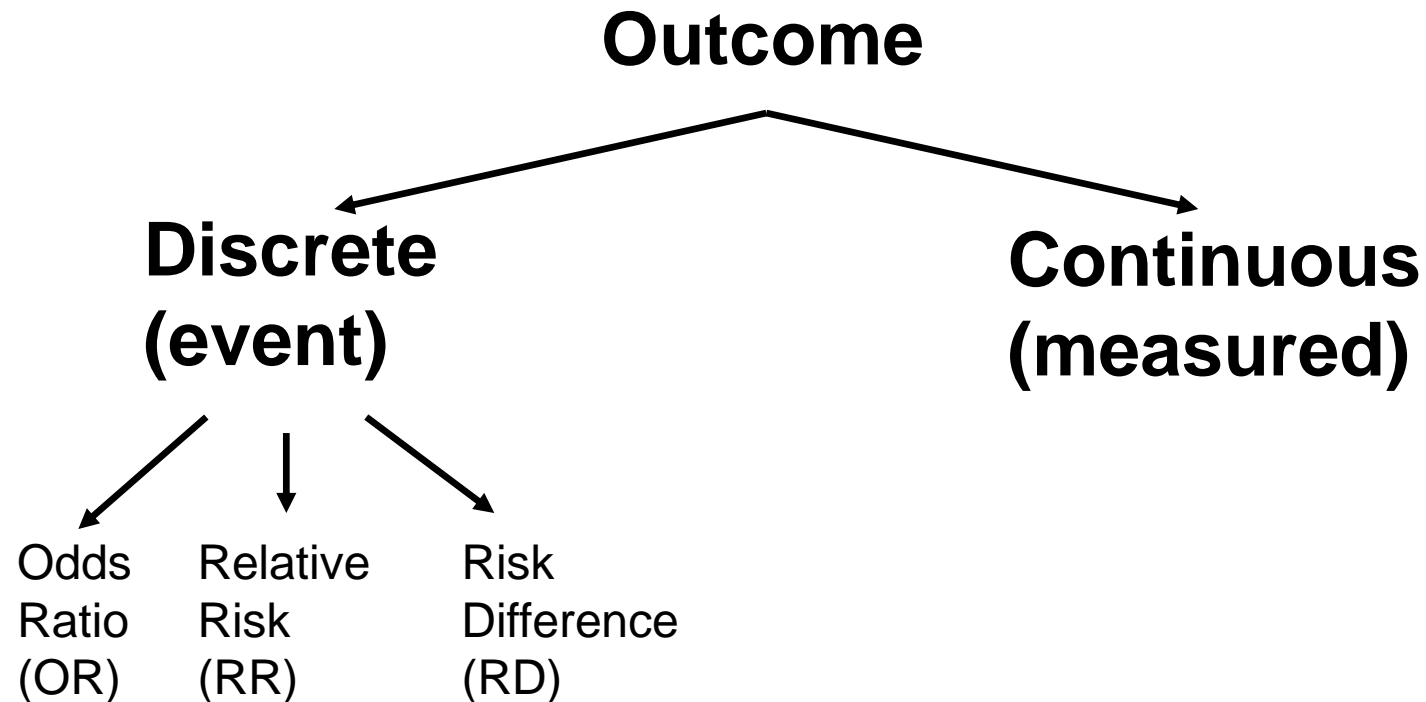
Interpretation of graphs and effect estimates in meta-analysis


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Forest Plot


- ◆ For each trial or study
 - estimate (square)
 - 95% confidence interval (CI) (line)
 - size (square) indicates weight allocated
- ◆ Solid vertical line of 'no effect'
 - if CI crosses line then effect not significant ($p > 0.05$)
- ◆ Horizontal axis
 - arithmetic: RD, MD, SMD
 - logarithmic: OR, RR
- ◆ Diamond represents combined estimate and 95% CI
- ◆ Dashed line plotted vertically through combined estimate

Effect Size Measures





What are dichotomous outcomes?

- ☐ when the outcome for every participant is one of two possibilities or events
 - alive or dead
 - healed or not healed
 - pregnant or not pregnant
- 

What were the chances of that?

■ **Risk** and **odds**

- express chance in numbers
- for dichotomous outcomes, express the chance within a group of being in one of two states
- particular statistical meanings, calculated differently

Risk

- 24 people drank coffee
6 developed a headache
- risk of a headache
 - = 6 headaches / 24 people who could have had one
 - = $6/24 = 1/4 = 0.25 = 25\%$

**risk = no. participants with event of interest
total no. participants**

Odds

- 24 people drank coffee
6 developed a headache
- odds of a headache
 - = 6 headaches/18 without headaches
 - = $6/18 = 1/3 = 0.33 = 1:3$ (not usually as %)

odds = $\frac{\text{no. participants with event of interest}}{\text{no. participants without event of interest}}$

Do risks and odds differ much?

- Two examples from caffeine trials

- 5 people with 'headaches' out of 65
- chance of having a headache

risk = $5/65 = 0.077$

odds = $5/60 = 0.083$

- 130 people 'still awake' out of 165
- chance of still being awake

risk = $130/165 = 0.79$

odds = $130/35 = 3.71$

Comparing two groups

	Headache	No headache	Total
Caffeine	17	51	68
Decaf	9	55	64
Total	26	106	132

Comparing two groups

- effect measures

- risk ratio (RR) (relative risk)
- odds ratio (OR)
- risk difference (RD) (absolute risk reduction)

- all estimates are uncertain, and should be presented with a confidence interval

Risk ratio

- risk of event with intervention
= **17/68=0.25**
- risk of event with control
= **9/64=0.14**
- risk ratio = $\frac{\text{intervention risk}}{\text{control risk}}$
- = $\frac{17/68}{9/64} = \frac{0.25}{0.14} = 1.79$
-

	Headache	No headache	Total
Caffeine	17	51	68
Decaf	9	55	64
Total	26	106	132

Where risk ratio = 1, there is no difference between the groups

Expressing it in words

- Risk ratio 1.79

- the risk of having a headache with treatment was 179% of the risk in the control group
- intervention increased the risk of headache by 79%

- **or for a reduction in risk:**

- Risk ratio 0.56

- the risk of having a headache with Decaf was 56% of the risk in the caffein group
- intervention reduced the risk of headache by 44%

Odds ratio

- odds of event with intervention
= **17/51**
- odds of event with control
= **9/55**
- odds ratio = $\frac{\text{intervention odds}}{\text{control odds}}$
- $\frac{17/51}{9/55} = 0.33 = 2.06$
- $\frac{9/55}{0.16}$
-

	Headache	No headache	Total
Caffeine	17	51	68
Decaf	9	55	64
Total	26	106	132

Where odds ratio = 1, there, is no difference between the groups

Expressing it in words

□ Odds ratio 2.06

- intervention doubled the odds of headache
- intervention increased the odds to 206% of the odds in the control group
- intervention increased the odds of headache by 106%

■ or for a reduction in odds:

□ Odds ratio 0.48

- Decaff reduced the odds of headache to 48% of the odds in the caffeine group
- Decaf reduced the odds of headache by 52%

Risk difference

	Headache	No headache	Total
Caffeine	17	51	68
Decaf	9	55	64
Total	26	106	132

- risk of event with intervention
= $17/68=0.25$
- risk of event with control
= $9/64=0.14$
- risk difference = risk with intervention – risk with control
 - $=17/68 - 9/64$
 - $= 0.25 - 0.14 = 0.11$
 - Where risk difference = 0, there is no difference between the groups

Expressing it in words

- Risk difference 0.11

- intervention increased the risk of headache by 11 percentage points
- 14 out of 100 people experienced a headache in the control group. 11 more people experienced a headache with caffeine.

- **or for a reduction in risk:**

- Risk difference -0.11

- intervention reduced the risk of headache by 11 percentage points
- 14 out of 100 people experienced a headache in the control group. 11 fewer people experienced a headache with caffeine.

Now it's your turn!

	Event	No Event	Total
Intervention	2	8	10
Control	5	5	10
Total	7	13	20

1. calculate:

- ☐ risk ratio for the effect of treatment on chance of event
- ☐ odds ratio for the effect of treatment on chance of event

2. express the results in words

The answers

- Risk ratio $= \frac{2/10}{5/10} = \frac{0.2}{0.5} = 0.4$

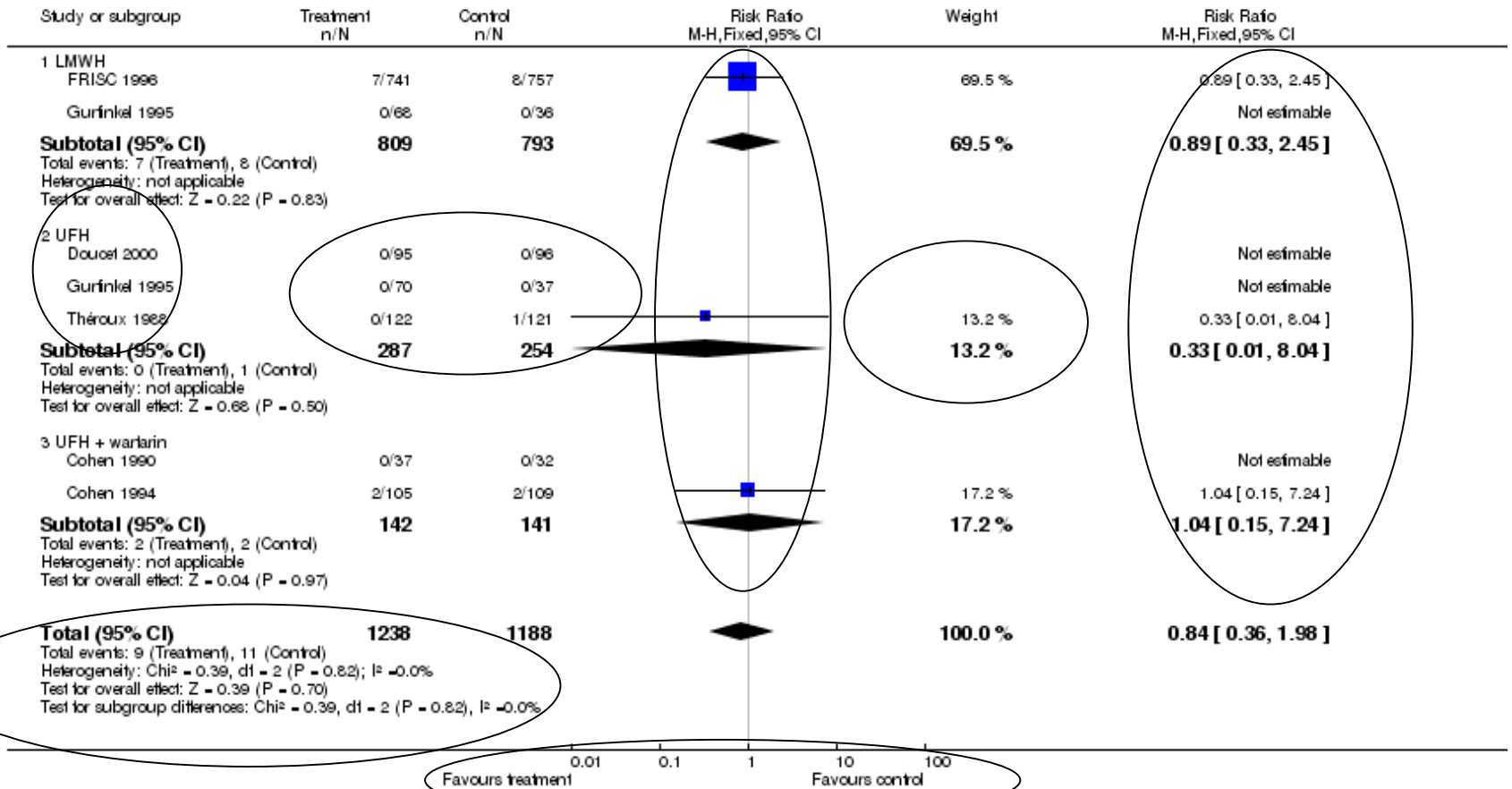
$$= \frac{2/8}{5/5} = \frac{0.25}{1} = 0.25$$

- Odds ratio

Communication

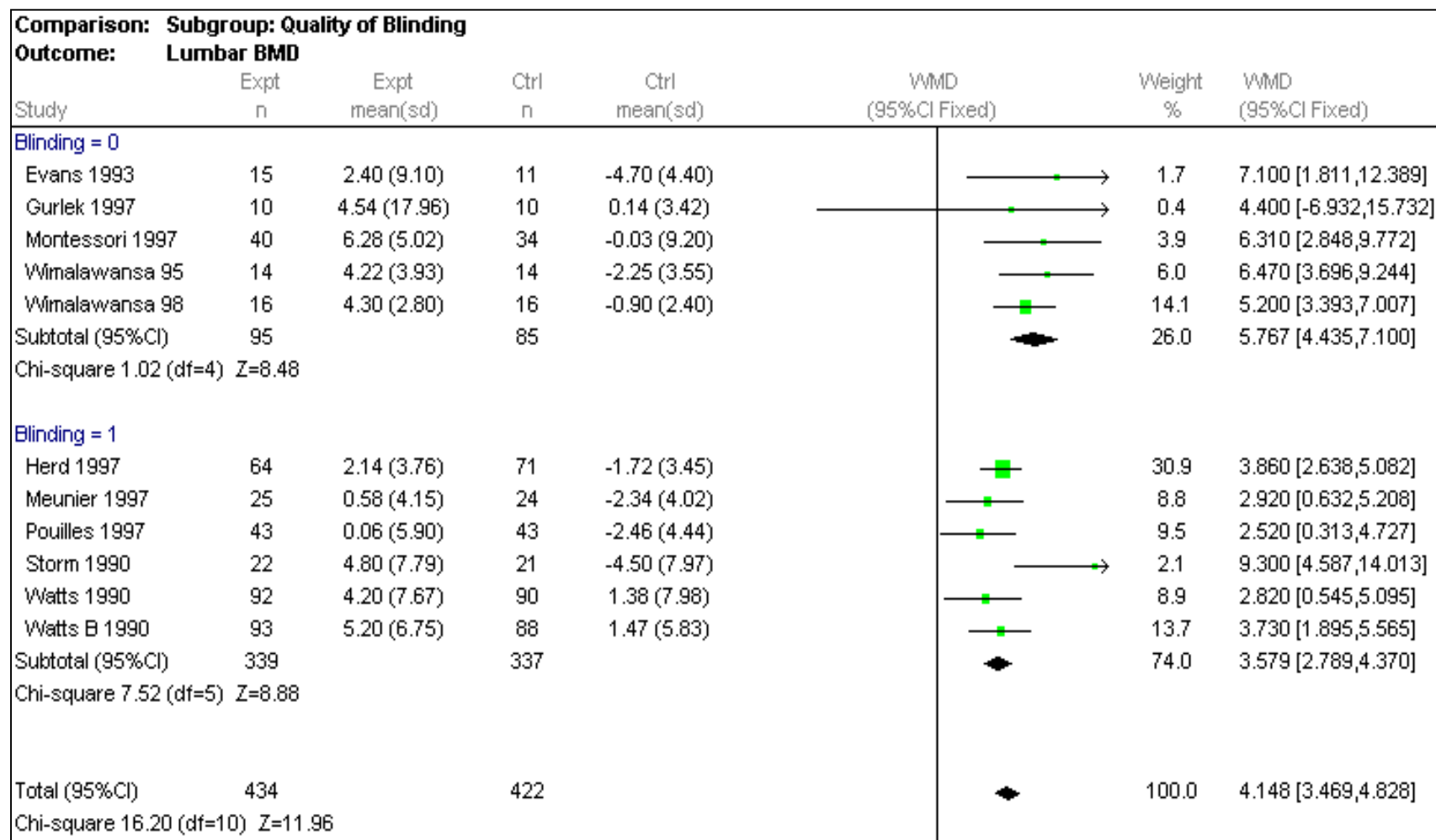
- OR is hard to understand, often misinterpreted
- RR is easier, but relative
 - can mean a very big or very small change
- RD is easiest
 - absolute measure of actual change in risk
 - easily converted to natural frequencies or NNT

Review: Heparin versus placebo for non-ST elevation acute coronary syndromes
 Comparison: 1 Incidence of death over all time periods
 Outcome: 1 Heparin vs placebo or untreated control



1.1. Comparison 1 Incidence of death over all time periods, Outcome 1 Heparin vs placebo or untreated control.

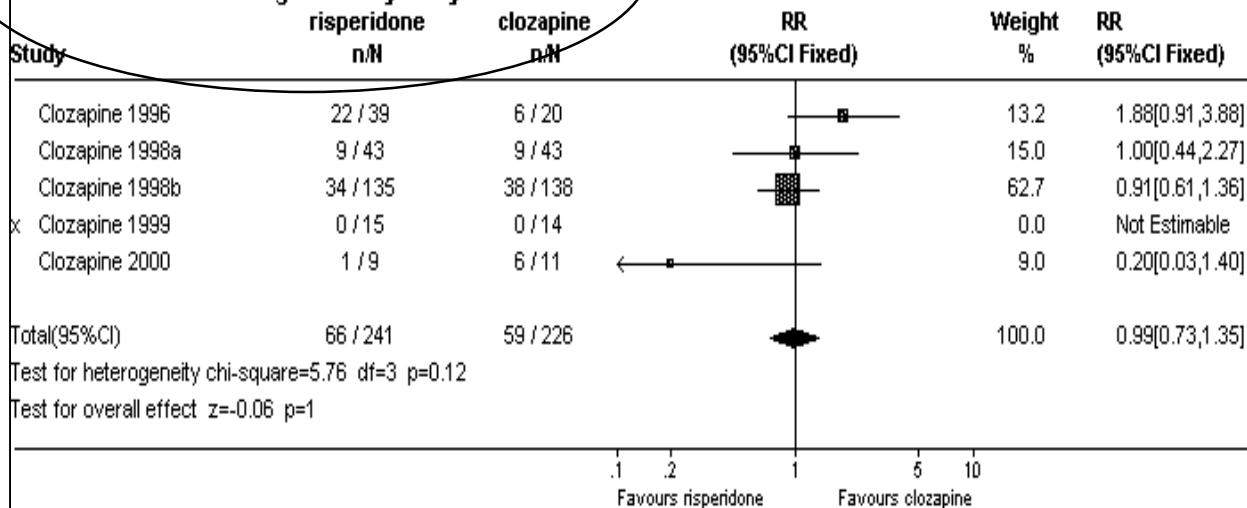
Forest plot



Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

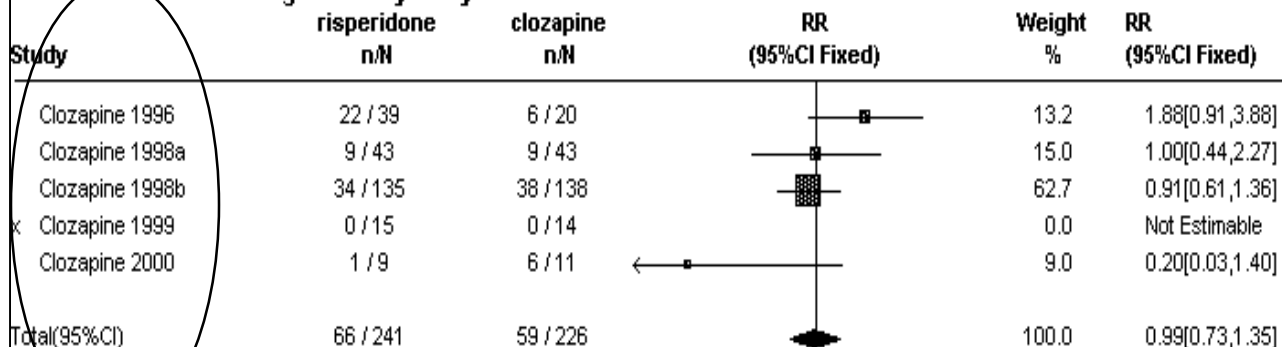
Outcome: 02 Leaving the study early



Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

Outcome: 02 Leaving the study early



Test for heterogeneity: chi-square=5.76 df=3 p=0.12

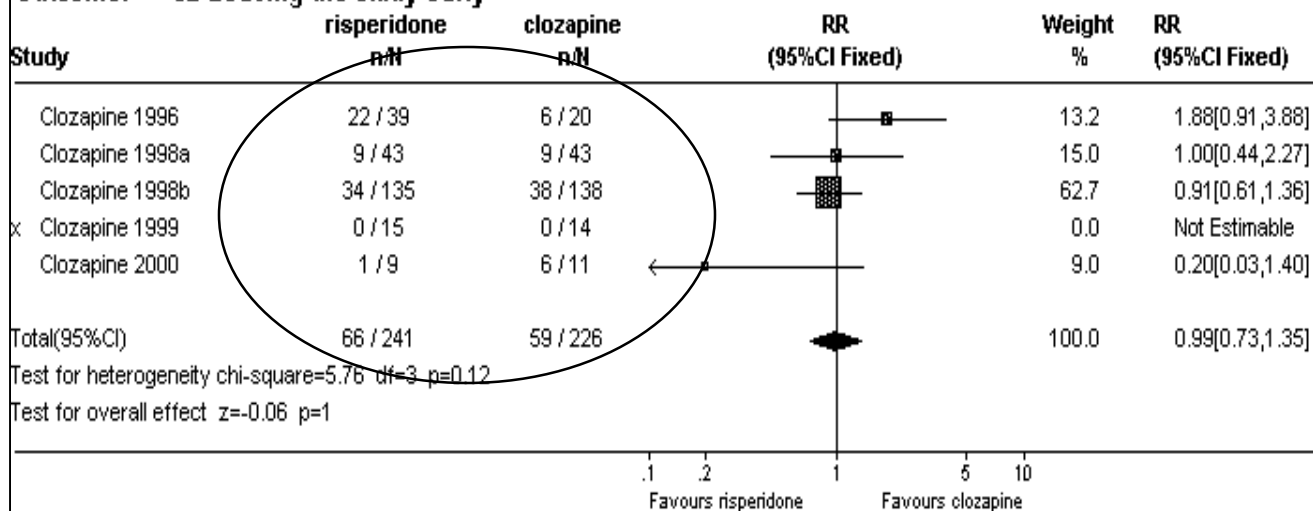
Test for overall effect: z=-0.06 p=1

.1 .2 1 5 10
Favours risperidone Favours clozapine

Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

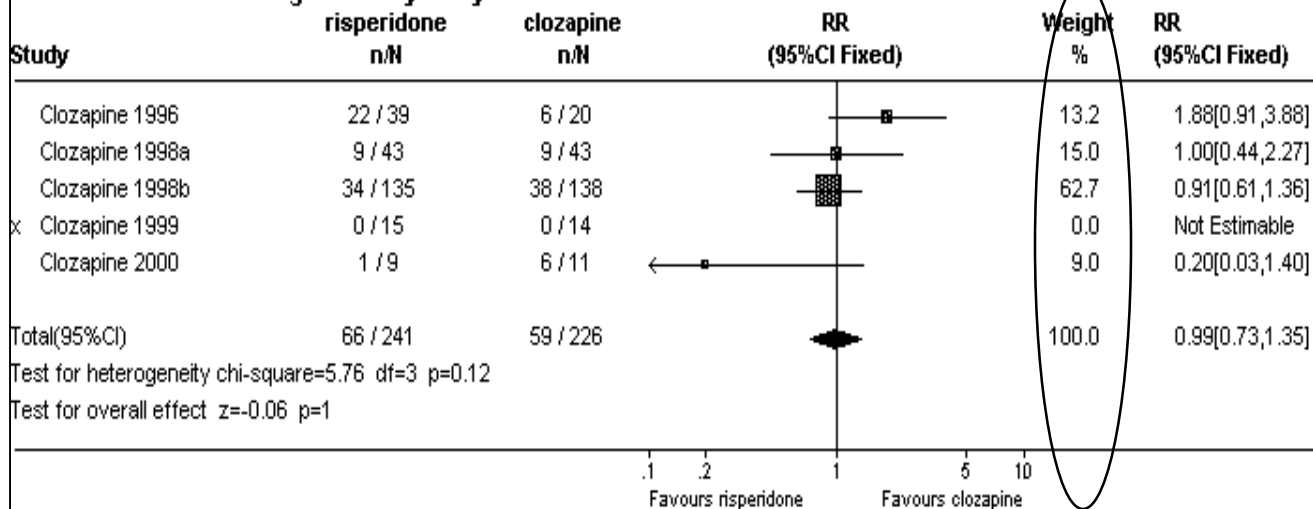
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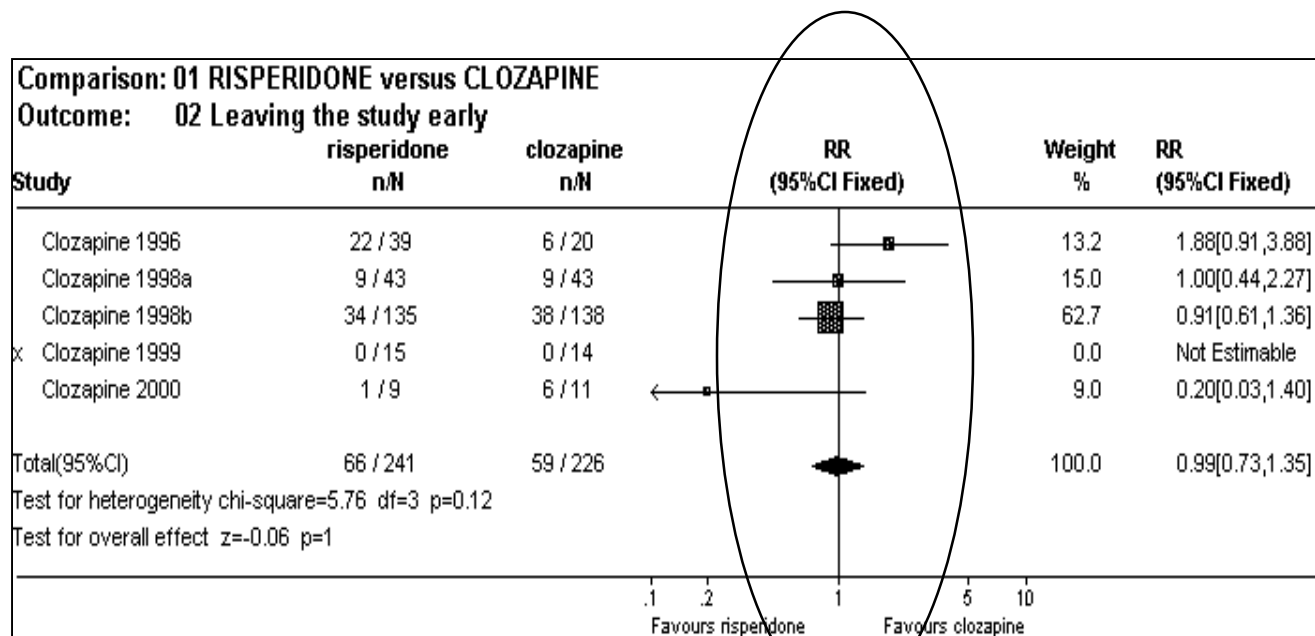
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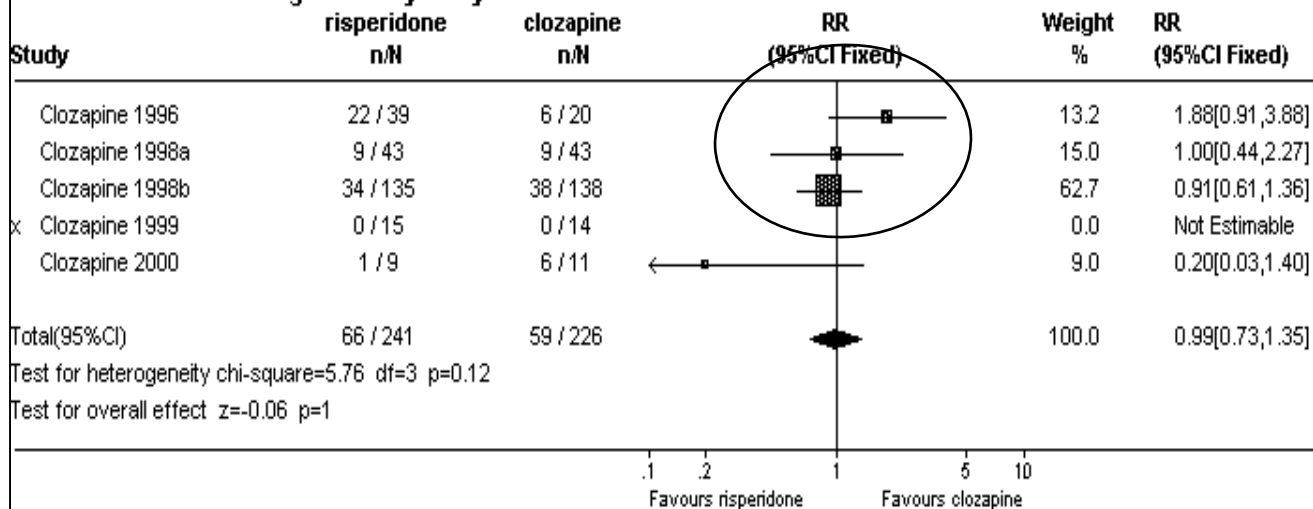
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Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

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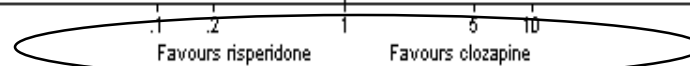
Comparison: 01 RISPERIDONE versus CLOZAPINE

Outcome: 02 Leaving the study early

Study	risperidone n/N	clozapine n/N	RR (95%CI Fixed)	Weight %	RR (95%CI Fixed)
Clozapine 1996	22 / 39	6 / 20		13.2	1.88[0.91,3.88]
Clozapine 1998a	9 / 43	9 / 43		15.0	1.00[0.44,2.27]
Clozapine 1998b	34 / 135	38 / 138		62.7	0.91[0.61,1.36]
Clozapine 1999	0 / 15	0 / 14		0.0	Not Estimable
Clozapine 2000	1 / 9	6 / 11		9.0	0.20[0.03,1.40]
Total(95%CI)	66 / 241	59 / 226		100.0	0.99[0.73,1.35]

Test for heterogeneity chi-square=5.76 df=3 p=0.12

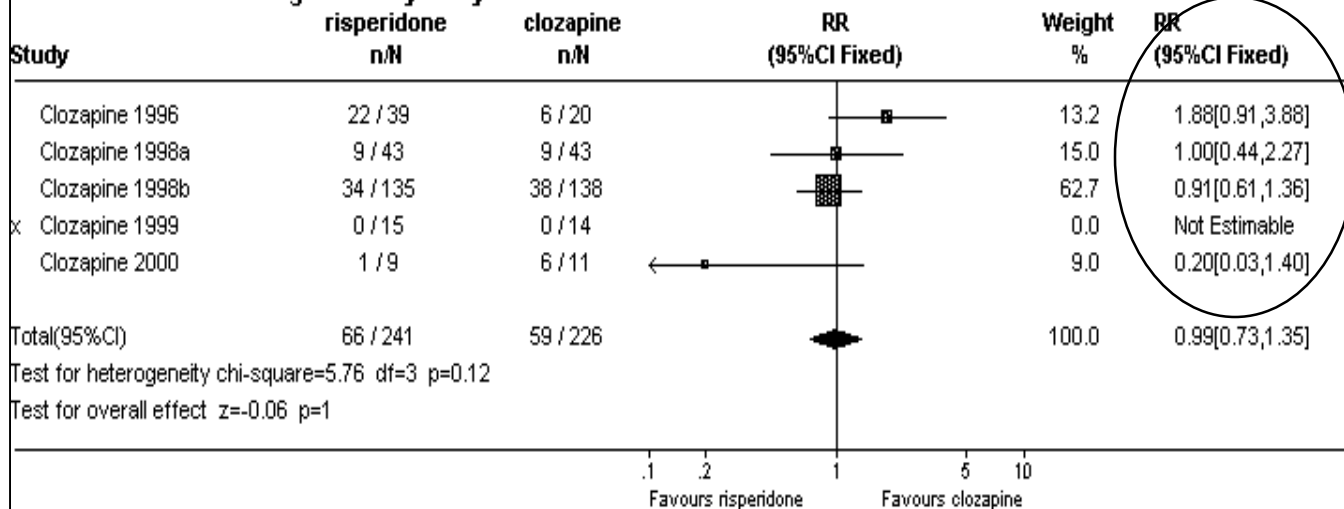
Test for overall effect z=-0.06 p=1



Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

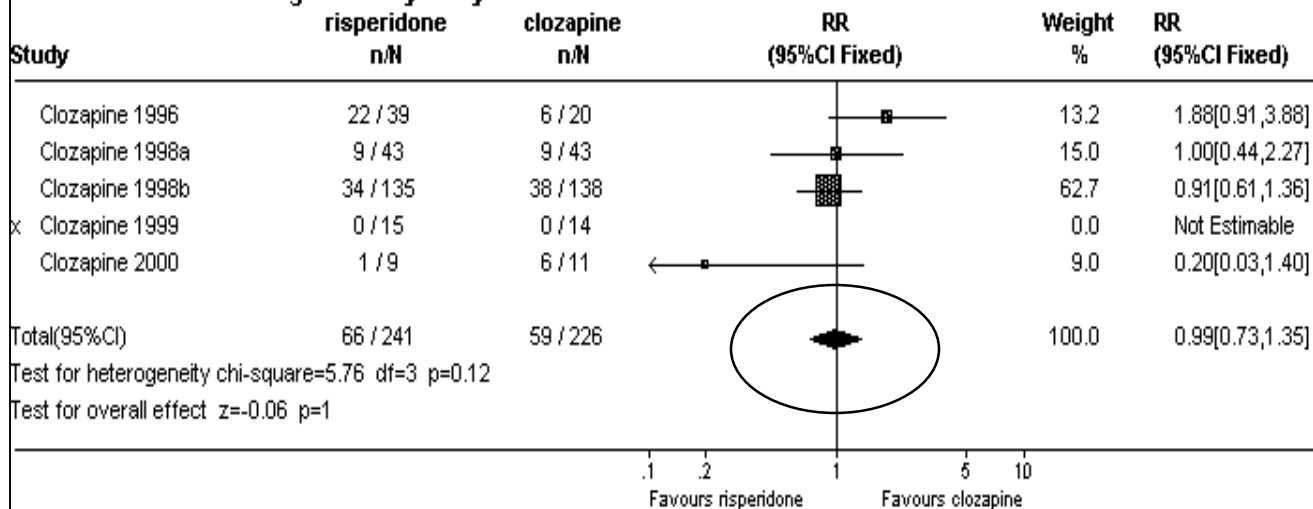
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Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

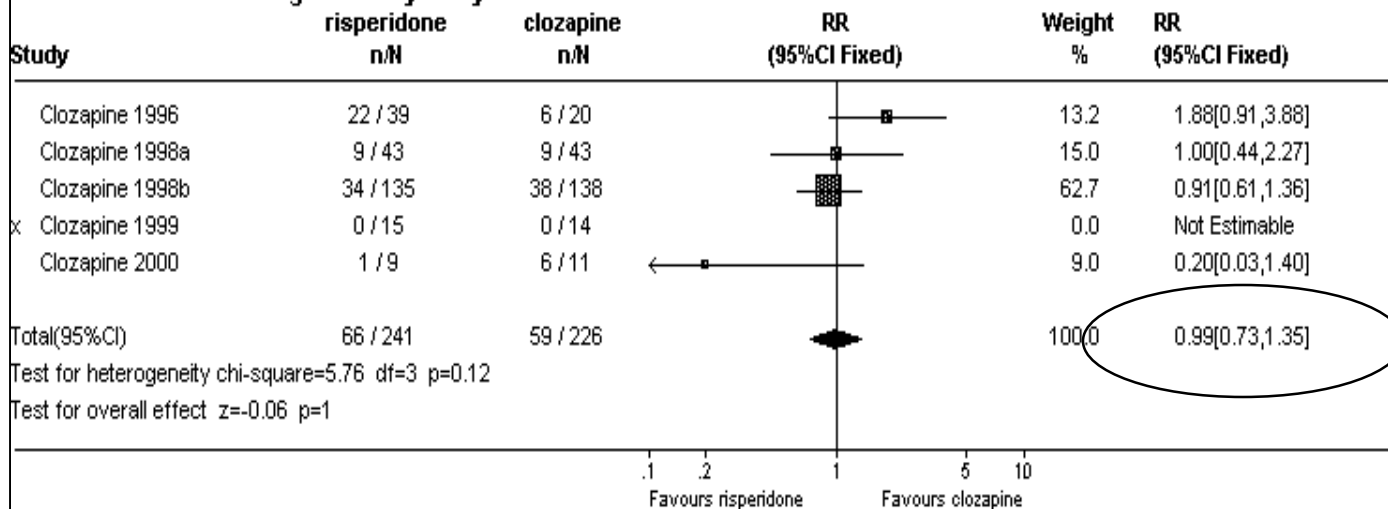
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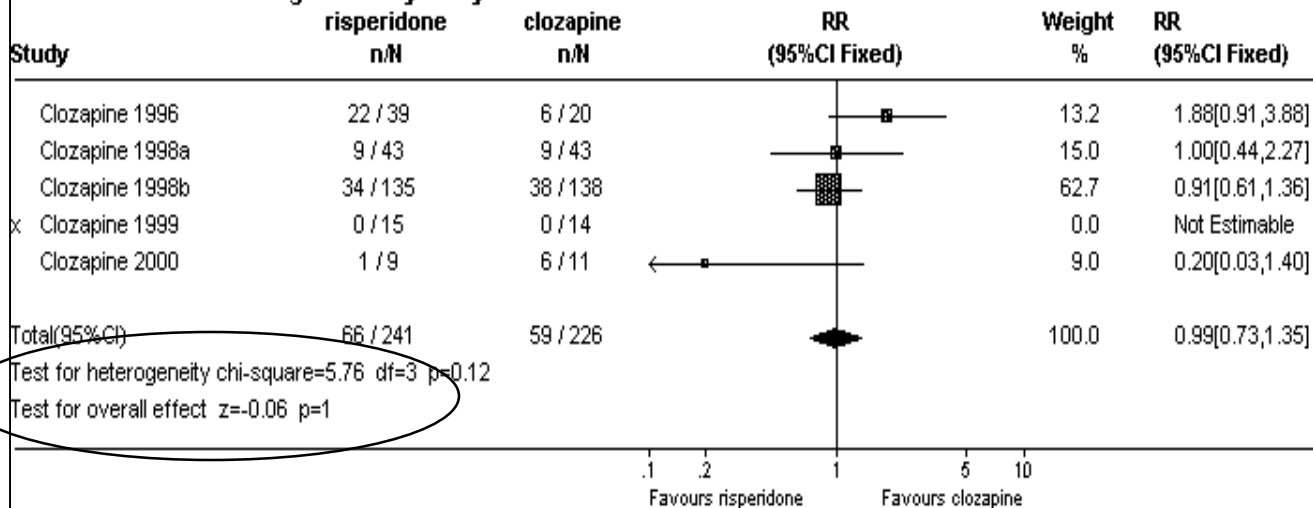
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Forest plot

Comparison: 01 RISPERIDONE versus CLOZAPINE

Outcome: 02 Leaving the study early





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