

**Cochrane** Database of Systematic Reviews

# Acupuncture for schizophrenia (Review)

Shen X, Xia J, Adams CE

Shen X, Xia J, Adams CE. Acupuncture for schizophrenia. *Cochrane Database of Systematic Reviews* 2014, Issue 10. Art. No.: CD005475. DOI: 10.1002/14651858.CD005475.pub2.

www.cochranelibrary.com

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



# TABLE OF CONTENTS

STRACT	•
AIN LANGUAGE SUMMARY	
MMARY OF FINDINGS	
CKGROUND	•
Figure 1	•
JECTIVES	
THODS	•
SULTS	
Figure 2	•
Figure 3	•
Figure 4	•
SCUSSION	••
THORS' CONCLUSIONS	•
KNOWLEDGEMENTS	•
	•
ARACTERISTICS OF STUDIES	•
TA AND ANALYSES	 r
ANTIPSYCHOTICS, Outcome 1 Global state: Not improved, endpoint.	E 
Analysis 1.2. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Mental state: 1a. General - average score (BPRS, endpoint, high score = worse).	Е •
Analysis 1.3. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Mental state: 1b. General - average score (BPRS, endpoint, high score = worse, medium-term) - subgroup analysis.	E -
Analysis 1.4. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Mental state: 2a. General - average score (PANSS, endpoint, high score = worse).	E
Analysis 1.5. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Mental state: 2b. General - average score (PANSS, endpoint, high score = worse, short-term) - subgroup analysis.	E -
Analysis 1.6. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Mental state: 2c. General - average score (PANSS, endpoint, high score = worse, short-term) - electroacupuncture subgroup analysis.	E -
Analysis 1.7. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Mental state: 2d. General - average score (PANSS, endpoint, high score = worse) - Skewed data.	E
Analysis 1.8. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 8 Mental state: 2e. General - not improved (PANSS), endpoint.	E 
Analysis 1.9. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 9 Mental state: 3. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse short-term) - Skewed data.	E 2,
Analysis 1.10. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 10 Mental state: 4a.Specific - average score - negative symptoms (SANS, endpoint, high score = worse).	E = 
Analysis 1.11. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 11 Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data.	E =
Analysis 1.12. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 12 Mental state: 5a. Specific - average score - depression (HAMD, endpoint, high score = worse short-term).	E 2,
Analysis 1.13. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 13 Mental state: 5b. Specific - not improved - depression (HAMD, reduced rate < 25%, short-term).	E
Analysis 1.14. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 14 Mental state: 7b. Specific - not improved - auditory hallucinations (PSYRAS-AH, reduction < 20% short-term)	Е 5,



Analysis 1.15. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 15 Mental state: 7a. Specific - average score - auditory hallucinations (PSYRAS-AH,endpoint, high score = worse, short-term).	100
Analysis 1.16. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 16 Mental state: 6. Specific - average score - depression (SDS, endpoint, high score = worse, short-term).	100
Analysis 1.17. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 17 Mental state: 8. Specific - not improved (auditory hallucinations, endpoint).	100
Analysis 1.18. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 18 Mental state: 9. Specific - average score - auditory hallucinations (SAHS, endpoint, high score = worse) - Skewed data.	101
Analysis 1.19. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 19 Mental state: 10. Specific - average score - hallucinations (BPRS [12th item], endpoint, high score = worse, short-term).	101
Analysis 1.20. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 20 Behaviour: Leaving the study early.	102
Analysis 1.21. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 21 Service outcomes: Time in hospital (days).	102
Analysis 1.22. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 22 Adverse effects: 1. General - average score (TESS, endpoint, high score = worse, short-term)	103
Analysis 1.23. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 23 Adverse effects: 2a. Specific - extrapyramidal symptoms.	103
Analysis 1.24. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 24 Adverse effects: 2b. Specific - extrapyramidal symptoms -overall (short-term) - subgroup analysis.	104
Analysis 1.25. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 25 Adverse effects: 3. Specific - Central Nervous System.	105
Analysis 1.26. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 26 Adverse effects: 4. Specific - anticholinergic symptoms.	106
Analysis 1.27. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 27 Adverse effects: 5. Specific - gastrointestinal system.	107
Analysis 1.28. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 28 Adverse effects: 6. Specific - cardiovascular symptoms (or headache).	108
Analysis 1.29. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 29 Adverse effects: 7a. Specific - metabolic system.	109
Analysis 1.30. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 30 Adverse effects: 7b. Specific - metabolic system - weight gain (short-term) - subgroup analysis.	110
Analysis 1.31. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS. Outcome 31 Adverse effects: 8. Specific - endocrine system.	110
Analysis 1.32. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS Outcome 32 Adverse effects: 9 Specific - lab test	111
Analysis 2.1. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 1 Global state: 1. Relapse (follow-up, long-term)	115
Analysis 2.2. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Global state: 2 Not improved (various similar criteria) endpoint (short-term)	115
Analysis 2.3. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Global state: 3a, CGL, average score - CGL-SL (endpoint, high score = worse, short-term)	116
Analysis 2.4. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Global state: 3b. CGL- average score - CGL-GL (endpoint, high score = worse, short-term) - Skewed data	116
Analysis 2.5. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Global state: 3c. CGL - average score - CGLEL (endpoint, high score = worse, short-term) - Skewed data	117
Analysis 2.6. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Mental state: 1a, General - average score (BPRS, endpoint, high score = worse)	117
Analysis 2.7. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Mental state: 1b. General- average score (BPPS, endpoint, high score = worse, short term), subgroup analysis	117
Analysis 2.8. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 8 Mental state: 1c. General - not improved (BPRS, short-term)	118
outcome o mental outcome a not improved (b) not official animalian an	

Copyright @ 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



Analysis 2.9. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 9 Mental state: 1d. General - average change scores (BPRS, low score = worse, short-term).	119
Analysis 2.10. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 10 Mental state: 2a. General - Average score (PANSS, endpoint, high score = worse, short-term) - Skew data Analysis 2.11. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS,	119 120
Outcome 11 Mental state: 2b. General - not improved (PANSS, reduced rate < 30%, short-term).	120
Outcome 12 Mental state: 3a. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term).	120
Analysis 2.13. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 13 Mental state: 3b. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data.	120
Analysis 2.14. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 14 Mental state: 4a. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term).	121
Analysis 2.15. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 15 Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data.	121
Analysis 2.16. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 16 Behaviour: Leaving the study early (short-term).	121
Analysis 2.17. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS,	122
Outcome 17 Adverse effects: 1a. General - average score (TESS, endpoint, high score = worse, short-term).	100
Outcome 18 Adverse effects: 1b. General - average score (TESS, endpoint, high score = worse, short-term) - Skewed data	122
Analysis 2.19. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS,	122
Outcome 19 Adverse effects: 2. Specific - average score (RESES, endpoint, high score = worse, short-term).	100
Outcome 20 Adverse effects: 3. Specific - extrapyramidal symptoms (short-term).	125
Analysis 2.21. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 21 Adverse effects: 4. Specific - Central Nervous System (short-term).	123
Analysis 2.22. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 22 Adverse effects: 5. Specific - anticholinergic symptoms (short-term).	124
Analysis 2.23. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 23 Adverse effects: 6. Specific - cardiovascular symptoms (short-term).	125
Analysis 2.24. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 24 Adverse effects: 7. Specific - skin infection (short-term).	126
Analysis 3.1. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 1 Global state: Not improved, endpoint (short-term).	127
Analysis 3.2. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Mental state: 1. General - average score (BPRS, endpoint, high score = worse, short-term).	127
Analysis 3.3. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Mental state: 2. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term).	128
Analysis 3.4. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Mental state: 3. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term).	128
Analysis 3.5. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Behaviour: Leaving the study early (short-term).	128
Analysis 3.6. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Adverse effects: 1. General - average scores (TESS, endpoint, short-term) - Skewed data.	129
Analysis 3.7. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Adverse effects: 2. Specific - extrapyramidal symptoms (short-term).	129
Analysis 3.8. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 8 Adverse effects: 3. Specific - numbness over the ear, upper extremity and chest on the treated side (short-term).	129
Analysis 4.1. Comparison 4 ACUPUNCTURE added to TCM DRUG versus TCM DRUG, Outcome 1 Global state: Not improved, endpoint (short-term).	130
Analysis 4.2. Comparison 4 ACUPUNCTURE added to TCM DRUG versus TCM DRUG, Outcome 2 Behaviour: Leaving the study early (short-term).	130
Analysis 5.1. Comparison 5 ACUPUNCTURE versus TCM DRUG, Outcome 1 Global state: Not improved, endpoint (short-term).	131
Analysis 5.2. Comparison 5 ACUPUNCTURE versus TCM DRUG, Outcome 2 Behaviour: Leaving the study early (short-term)	132

Copyright @ 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



Analysis 6.1. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 1 Behaviour: Leaving the study early (short-term).	132
Analysis 6.2. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 2 Adverse effects: 1. Specific - back pain (short-term).	133
Analysis 6.3. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 3 Adverse effects: 2. Spinal fracture (short-term).	133
ADDITIONAL TABLES	133
APPENDICES	141
WHAT'S NEW	145
HISTORY	145
CONTRIBUTIONS OF AUTHORS	145
DECLARATIONS OF INTEREST	146
SOURCES OF SUPPORT	146
DIFFERENCES BETWEEN PROTOCOL AND REVIEW	146
INDEX TERMS	146



# Acupuncture for schizophrenia

Xiaohong Shen<sup>1</sup>, Jun Xia<sup>2</sup>, Clive E Adams<sup>2</sup>

<sup>1</sup>Shanghai Shuguang Hospital, Shanghai University of Traditional Chinese Medicine, Shanghai, China. <sup>2</sup>Cochrane Schizophrenia Group, The University of Nottingham, Nottingham, UK

**Contact:** Xiaohong Shen, Shanghai Shuguang Hospital, Shanghai University of Traditional Chinese Medicine, No 185 Pu'an Road, Shanghai, 200021, China. stalashen@hotmail.com.

**Editorial group:** Cochrane Schizophrenia Group. **Publication status and date:** New search for studies and content updated (no change to conclusions), published in Issue 10, 2014.

**Citation:** Shen X, Xia J, Adams CE. Acupuncture for schizophrenia. *Cochrane Database of Systematic Reviews* 2014, Issue 10. Art. No.: CD005475. DOI: 10.1002/14651858.CD005475.pub2.

Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

# ABSTRACT

# Background

Acupuncture, with many categories such as traditional acupuncture, electroacupuncture, laser acupuncture, and acupoint injection, has been shown to be relatively safe with few adverse effects. It is accessible and inexpensive, at least in China, and is likely to be widely used there for psychotic symptoms.

#### Objectives

To review the effects of acupuncture, alone or in combination treatments compared with placebo (or no treatment) or any other treatments for people with schizophrenia or related psychoses.

# Search methods

We searched Cochrane Schizophrenia Group's Trials Register (February 2012), which is based on regular searches of CINAHL, BIOSIS, AMED, EMBASE, PubMed, MEDLINE, PsycINFO and clinical trials registries. We also inspected references of identified studies and contacted relevant authors for additional information.

#### **Selection criteria**

We included all relevant randomised controlled trials involving people with schizophrenia-like illnesses, comparing acupuncture added to standard dose antipsychotics with standard dose antipsychotics alone, acupuncture added to low dose antipsychotics with standard dose antipsychotics, acupuncture with antipsychotics, acupuncture added to Traditional Chinese Medicine (TCM) drug with TCM drug, acupuncture with TCM drug, electric acupuncture convulsive therapy with electroconvulsive therapy.

#### Data collection and analysis

We reliably extracted data from all included studies, discussed any disagreement, documented decisions and contacted authors of studies when necessary. We analysed binary outcomes using a standard estimation of risk ratio (RR) and its 95% confidence interval (CI). For continuous data, we calculated mean differences with 95% CI. For homogeneous data we used fixed-effect model. We assessed risk of bias for included studies and created 'Summary of findings' tables using GRADE.

# **Main results**

After an update search in 2012 the review now includes 30 studies testing different forms of acupuncture across six different comparisons. All studies were at moderate risk of bias.

When acupuncture plus standard antipsychotic treatment was compared with standard antipsychotic treatment alone, people were at less risk of being 'not improved' (n = 244, 3 RCTs, medium-term RR 0.40 CI 0.28 to 0.57, *very low quality evidence*). Mental state findings were



mostly consistent with this finding as was time in hospital (n = 120, 1 RCT, days MD -16.00 CI -19.54 to -12.46, *moderate quality evidence*). If anything, adverse effects were less for the acupuncture group (e.g. central nervous system, insomnia, short-term, n = 202, 3 RCTs, RR 0.30 CI 0.11 to 0.83, *low quality evidence*).

When acupuncture was added to low dose antipsychotics and this was compared with standard dose antipsychotic drugs, relapse was less in the experimental group (n = 170, 1 RCT, long-term RR 0.57 CI 0.37 to 0.89, *very low quality evidence*) but there was no difference for the outcome of 'not improved'. Again, mental state findings were mostly consistent with the latter. Incidences of extrapyramidal symptoms akathisia, were less for those in the acupuncture added to low dose antipsychotics group (n = 180, 1 RCT, short-term RR 0.03 CI 0.00 to 0.49, *low quality evidence*) - as dry mouth, blurred vision and tachycardia.

When acupuncture was compared with antipsychotic drugs of known efficacy in standard doses, there were equivocal data for outcomes such as 'not improved' using different global state criteria. Traditional acupuncture added to TCM drug had benefit over use of TCM drug alone (n = 360, 2 RCTs, RR no clinically important change 0.11 Cl 0.02 to 0.59, *low quality evidence*), but when traditional acupuncture was compared with TCM drug directly there was no significant difference in the short-term. However, we found that participants given electroacupuncture were significantly less likely to experience a worsening in global state (n = 88, 1 RCT, short-term RR 0.52 Cl 0.34 to 0.80, *low quality evidence*).

In the one study that compared electric acupuncture convulsive therapy with electroconvulsive therapy there were significantly different rates of spinal fracture between the groups (n = 68, 1 RCT, short-term RR 0.33 Cl 0.14 to 0.81, *low quality evidence*). Attrition in all studies was minimal. No studies reported death, engagement with services, satisfaction with treatment, quality of life, or economic outcomes.

# **Authors' conclusions**

Limited evidence suggests that acupuncture may have some antipsychotic effects as measured on global and mental state with few adverse effects. Better designed large studies are needed to fully and fairly test the effects of acupuncture for people with schizophrenia.

# PLAIN LANGUAGE SUMMARY

# Acupuncture for schizophrenia

Although acupuncture or Traditional Chinese Medicine has been practised for over 2000 years in China and the Far East, especially in Korea and Japan, it is a relatively new form of treament for physical and psychological conditions in the West. Acupuncture inserts needles into the skin to stimulate specific points of the body (acupoints). The aim is to achieve balance and harmony of the body.

Schizophrenia is a serious mental illness and is usually treated using antipsychotic medication. However, although effective, antipsychotic medication can cause side-effects (such as sleepiness, weight gain and even dribbling). Acupuncture has been shown to have very few negative effects on the individual and could be more socially acceptable and tolerable for people with mental health problems. Acupuncture may also be less expensive than drugs made by pharmaceutical companies, so reducing costs to individuals and health services.

This reviews looks at the effectiveness of various types of acupuncture as treatment for people with schizophrenia. An update search for studies was carried out in 2012 and found 30 studies that randomised participants who were receiving antipsychotic medication to receive additional acupuncture or standard care.

Although some of the studies did favour acupuncture when combined with antipsychotics, the information available was small scale and rated to be very low or low quality by the review authors, so not completely provable and valid. Depression was reduced when combining acupuncture with antipsychotic medication, but again this finding came from small-scale research, so cannot be clearly shown to be true. The review concludes that people with mental health problems, policy makers and health professionals need much better evidence in order to establish if there are any potential benefits to acupuncture.

This means that the question of whether acupuncture is of benefit to people, and whether it is of greater benefit than antipsychotic medication, remains unanswered. There is not enough information to establish that acupuncture is of benefit or harm to people with mental health problems.

Benjamin Gray, Service User and Service User Expert, Rethink Mental Illness.

# SUMMARY OF FINDINGS

Summary of findings for the main comparison. ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia

ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia

Patient or population: patients with schizophrenia

Intervention: ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	No of Partici- pants	Quality of the evidence	Comments	
	Assumed risk	Corresponding risk		(studies)	(GRADE)		
	Control	ACUPUNCTURE added to STANDARD DOSE AN- TIPSYCHOTICS versus STANDARD DOSE AN- TIPSYCHOTICS					
Global state: Not im- proved, endpoint - medi- um-term (various similar criteria)	557 per 1000	<b>223 per 1000</b> (156 to 318)	<b>RR 0.4</b> (0.28 to 0.57)	244 (3 study)	⊕⊝⊝⊝ very low <sup>1,2</sup>	Relapse was not reported but 'no clinically important change in glob- al state' was reported and another outcome without clear duration in- dicated no difference between two comparison groups.	
Mental state: PANSS (not improved, reduced rated < 25%, short-term)	432 per 1000	<b>281 per 1000</b> (194 to 406)	<b>RR 0.65</b> (0.45 to 0.94)	197 (3 studies)	⊕000 very low <sup>1,2</sup>	Data from PANSS were equivocal because of different criteria. Other mental state findings were mostly consistent with this finding.	
Behaviour: Leaving the study early (short-term)	7 per 1000	<b>9 per 1000</b> (2 to 38)	<b>RR 1.33</b> (0.33 to 5.45)	870 (10 studies)	⊕000 very low <sup>2,3,4</sup>	Similar outcomes at medium-term.	
Service outcomes: Time in hospital (days)		The mean service out- comes: time in hospital (days) in the interven- tion groups was <b>16 lower</b> (19.54 to 12.46 lower)		120 (1 study)	⊕⊕⊕⊝ moderate <sup>2</sup>	Only one study reported service outcomes - time in hospital.	

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Adverse effects: Central Nervous System - insom- nia (short-term)	131 per 1000	<b>39 per 1000</b> (14 to 109)	<b>RR 0.30</b> (0.11 to 0.83)	202 (3 studies)	⊕⊕⊙⊙ low <sup>2,3</sup>	Only insomnia rate indicated dif- ference between two compared groups.		
Quality of life: No clinically important change in quali- ty of life	See comment	See comment	Not estimable	-	See comment	Not reported.		
Economic outcomes: Cost of care	See comment	See comment	Not estimable	-	See comment	Not reported.		
*The basis for the <b>assumed ris</b> based on the assumed risk in t <b>CI:</b> Confidence interval; <b>RR:</b> Ri	<b>sk</b> (e.g. the mediar the comparison gro isk ratio;	control group risk across stu oup and the <b>relative effect</b> o	udies) is provided in f the intervention (	n footnotes. The <b>co</b> and its 95% CI).	rresponding risk (a	nd its 95% confidence interval) is		
GRADE Working Group grades High quality: Further research Moderate quality: Further res Low quality: Further research Very low quality: We are very	of evidence h is very unlikely to search is likely to h i is very likely to ha uncertain about tl	o change our confidence in th ave an important impact on o ve an important impact on o ne estimate.	e estimate of effectour confidence in t ur confidence in th	t. he estimate of effec e estimate of effect	t and may change t and is likely to cha	he estimate. nge the estimate.		
<ul> <li><sup>1</sup> Risk of bias: rated 'very serious' - Unblinding of participants and personnel and incomplete outcome data.</li> <li><sup>2</sup> Imprecision: rated 'serious' - Few participants and/or few events.</li> <li><sup>3</sup> Risk of bias: rated 'serious' - Unblinding of participants and personnel.</li> <li><sup>4</sup> Publication bias: rated 'strong suspected' - One author worked for drug industry.</li> </ul>								
Summary of findings 2. AC	UPUNCTURE ad	ded to LOW DOSE ANTIP	SYCHOTICS vers	us STANDARD DC	SE ANTIPSYCHO	TICS for schizophrenia		

# ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia

**Patient or population:** patients with schizophrenia **Intervention:** ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Outcomes	Illustrative comparative risks* (95% CI)	Relative effect (95% CI)	No of Partici- pants (studies)	Quality of the evidence (GRADE)	Comments
•	Assumed risk Corresponding risk Control ACUPUNCTURE added to LOW DOSE ANTIPSY- CHOTICS ver-	-	(studies)		

4

Cochrane Library

Trusted evidence. Informed decisions. Better health.

		sus STANDARD DOSE ANTIPSY- CHOTICS				
Global state: Relapse (follow-up, long-term)	431 per 1000	<b>246 per 1000</b> (159 to 383)	<b>RR 0.57</b> (0.37 to 0.89)	170 (1 study)	⊕⊙⊙⊙ very low <sup>1,2,3</sup>	Only one study reported this outcome. Oth- er global state findings reported no difference between the two comparison groups.
Mental state: BPRS (not improved, reduced rate < 30%, short-term)	133 per 1000	<b>100 per 1000</b> (24 to 409)	<b>RR 0.75</b> (0.18 to 3.07)	60 (1 study)	⊕⊙⊙⊝ very low <sup>4,5</sup>	Though measured with different criteria of 'no clinically important change in general mental state', there was no difference be- tween the two comparison groups at short- term and mostly similar results from other mental state findings.
Behaviour: Leaving the study early (short- term)	19 per 1000	<b>16 per 1000</b> (6 to 45)	<b>RR 0.81</b> (0.29 to 2.29)	662 (8 studies)	⊕000 very low <sup>2,3,4</sup>	
Service outcomes: Hos- pitalisation	See comment	See comment	Not estimable	-	See comment	Not reported.
Adverse effects: Ex- trapyramidal symp- toms - specific - akathisia (short-term)	185 per 1000	<b>6 per 1000</b> (0 to 91)	<b>RR 0.03</b> (0 to 0.49)	180 (1 study)	⊕⊕⊝⊝ low <sup>2,4</sup>	Similar to the akathisia data, acupuncture added to low dose antipsychotics reduced participants experiencing dry mouth, blurred vision, tachycardia at short-term. Only one study focused on acupoint catgut treatment relative adverse effects but did not find skin infection in either groups.
Quality of life: No clini- cally important change in quality of life	See comment	See comment	Not estimable	-	See comment	Not reported.
Economic outcomes: Cost of care	See comment	See comment	Not estimable	-	See comment	Not reported.
*The basis for the <b>assume</b> based on the assumed risk	<b>d risk</b> (e.g. the means a in the comparisor	dian control group risl	k across studies) is <b>ve effect</b> of the int	provided in footno	otes. The <b>correspond</b> 95% Cl).	<b>ling risk</b> (and its 95% confidence interval) is

Cl: Confidence interval; RR: Risk ratio;

GRADE Working Group grades of evidence

High quality: Further research is very unlikely to change our confidence in the estimate of effect.

**Moderate quality:** Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate. **Low quality:** Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Cochrane Library

$\sim$	-
-	Ξ.
ω.	=
<u> </u>	5
7	÷++
0	-
9	
N	Ð
0	
ř	2
4	- 21
-	S
Ч	
Ð	≝.
0	N
ò	0
ŏ	σ
<u> </u>	5
5	
ē	œ
7	2
Ð	α.
0	-
0	T
Ě	~
a	2
Ъ	⊇.
ō	D.
5	5
Ē	2
<u> </u>	
0	
п	
·	
Ψ	
<u> </u>	
ъ	
_	
S	
sh	
ishe	
shed	
shed b	
ished by	
ished by J	
ished by Jo	
ished by Joh	
ished by Johr	
ished by John \	
ished by John W	
ished by John Wil	
ished by John Wile	
ished by John Wiley	
ished by John Wiley &	
shed by John Wiley &	
ished by John Wiley & S	
ished by John Wiley & Soi	
ished by John Wiley & Son:	
ished by John Wiley & Sons,	
ished by John Wiley & Sons, I	
ished by John Wiley & Sons, Lt	

Cop

# Very low quality: We are very uncertain about the estimate.

<sup>1</sup> Risk of bias: rated 'very serious' - Unblinding of participants and personnel and incomplete outcome data.

<sup>2</sup> Imprecision: rated 'serious' - Few participants and/or few events.

<sup>3</sup> Publication bias: rated 'strongly suspected' - There was some difference between two references of the same study.

<sup>4</sup> Risk of bias: rated 'serious' - Unblinding of participants and personnel.

<sup>5</sup> Imprecision: rated 'very serious' - Few participants and/or few events and wide confidence intervals.

# Summary of findings 3. ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia

# ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia

Patient or population: patients with schizophrenia Intervention: ACUPUNCTURE versus ANTIPSYCHOTICS

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	No of Partici- pants (studies)	Quality of the evidence (GRADE)	Comments
	Assumed risk	Corresponding risk		(otuarco)	(01012-2)	
	Control	ACUPUNCTURE versus ANTIPSY- CHOTICS				
Global state: Not im- proved (talk, behaviour and expression still ex- isted as before; the re- duction of BPRS < 29%), endpoint (short-term)	317 per 1000	<b>104 per 1000</b> (60 to 187)	<b>RR 0.33</b> (0.19 to 0.59)	240 (2 studies)	⊕⊝⊝⊝ very low <sup>1,2</sup>	Relapse was not reported but 'no clini- cally important change in global state' was reported and another three out- comes with different but similar criteria indicated no difference between the two comparison groups in the short-term.
Mental state: BPRS, endpoint (high score = worse, short-term) - tra- ditional acupuncture		The mean mental state: BPRS, end- point (high score = worse, short-term) - traditional acupunc- ture in the interven- tion groups was <b>11.56 lower</b> (16.36 to 6.76 lower)		32 (1 study)	⊕⊙⊙ <b>very low</b> <sup>1,2</sup>	The outcome of 'no clinically important change in general mental state' was not reported but average endpoint BPRS da- ta were reported as was the SAPS score. There was no difference between the two comparison groups in the short- term using either score.
Behaviour: Leaving the study early (short-term)	See comment	See comment	Not estimable	421 (6 studies)	⊕⊕©© low <sup>1,3</sup>	No participants left each compared group early.



pitalisation	See comment	See comment	Not estimable	-	See comment	Not reported.		
Adverse effects: Ex- trapyramidal symptoms (short-term)	800 per 1000	<b>40 per 1000</b> (0 to 664)	<b>RR 0.05</b> (0 to 0.83)	21 (1 study)	⊕⊕⊙⊙ low <sup>1,3</sup>	One study reported laser acupuncture relative adverse effects - numbness over ear, upper extremity and chest on the treated side, but no difference between the two comparison groups.		
Quality of life: No clini- cally important change in quality of life	See comment	See comment	Not estimable	-	See comment	Not reported.		
Economic outcomes: Cost of care	See comment	See comment	Not estimable	-	See comment	Not reported.		
*The basis for the <b>assumed risk</b> (e.g. the median control group risk across studies) is provided in footnotes. The <b>corresponding risk</b> (and its 95% confidence interval) is based on the assumed risk in the comparison group and the <b>relative effect</b> of the intervention (and its 95% CI).								
<b>Cl:</b> Confidence interval; <b>RR:</b> GRADE Working Group grad <b>High quality:</b> Further resea	Risk ratio; es of evidence rch is very unlikely	to change our confidence	in the estimate of	effect.	offect and may she	ngo the estimate		
CI: Confidence interval; RR: GRADE Working Group grad High quality: Further resea Moderate quality: Further Low quality: Further resea Very low quality: We are ve Risk of bias: rated 'serious' - Imprecision: rated 'very serious'	es of evidence rch is very unlikely research is likely to rch is very likely to f ery uncertain about - Unblinding of part ious' - Few participa - Few participants a	to change our confidence have an important impact have an important impact the estimate. icipants and personnel. ants and/or few events and and/or few events.	in the estimate of a t on our confidence on our confidence d wide confidence i	effect. e in the estimate of e in the estimate of e ntervals.	effect and may cha ffect and is likely t	inge the estimate. o change the estimate.		
CI: Confidence interval; RR: GRADE Working Group grad High quality: Further resea Moderate quality: Further Low quality: Further resea Very low quality: We are ver Risk of bias: rated 'serious' Imprecision: rated 'very serious' Imprecision: rated 'serious'	es of evidence rch is very unlikely research is likely to rch is very likely to h ery uncertain about Unblinding of part ious' - Few participa - Few participants a ACUPUNCTURE a	to change our confidence have an important impact nave an important impact the estimate. icipants and personnel. ants and/or few events and and/or few events.	in the estimate of a t on our confidence on our confidence d wide confidence i	effect. in the estimate of in the estimate of e ntervals. <b>For schizophreni</b>	effect and may cha ffect and is likely t	inge the estimate. o change the estimate.		
CI: Confidence interval; RR: GRADE Working Group grad High quality: Further resea Moderate quality: Further Low quality: Further resea Very low quality: We are ver Risk of bias: rated 'serious' Imprecision: rated 'very series Imprecision: rated 'serious' Cummary of findings 4. ACUPUNCTURE plus TCM E Patient or population: pat Intervention: ACUPUNCTU	Risk ratio; es of evidence rch is very unlikely research is likely to rch is very likely to h ery uncertain about Unblinding of part ious' - Few participa - Few participants a ACUPUNCTURE a DRUG versus TCM D ients with schizoph RE added to TCM D	to change our confidence have an important impact the estimate. icipants and personnel. ants and/or few events and and/or few events. added to TCM DRUG ver PRUG for schizophrenia renia RUG versus TCM DRUG	in the estimate of o t on our confidence on our confidence d wide confidence i	effect. In the estimate of e in the estimate of e ntervals.	effect and may cha ffect and is likely t	inge the estimate.		
CI: Confidence interval; RR: GRADE Working Group grad High quality: Further resea Moderate quality: Further Low quality: Further resea Very low quality: We are ver Risk of bias: rated 'serious' - Imprecision: rated 'very seri Imprecision: rated 'serious' Summary of findings 4. ACUPUNCTURE plus TCM I Patient or population: pat Intervention: ACUPUNCTU	Risk ratio; es of evidence rch is very unlikely to rch is very likely to f ery uncertain about Unblinding of part ious' - Few participa - Few participants a ACUPUNCTURE a DRUG versus TCM D ients with schizoph RE added to TCM D Illustrativ (95% CI)	to change our confidence have an important impact have an important impact the estimate. icipants and personnel. ants and/or few events and and/or few events. and/or few events.	in the estimate of o ton our confidence on our confidence d wide confidence i rsus TCM DRUG Relative effect (95% CI)	effect. e in the estimate of in the estimate of e ntervals. For schizophreni	effect and may cha iffect and is likely t a Quality of the evidence	o change the estimate.		

7



Trusted evidence. Informed decisions. Better health.

	Control	ACUPUNCTURE added to TCM DRUG versus TCM DRUG				
Global state: Not improved (talk, behaviour and expression still existed as before, the re- duction of BPRS < 29%), end- point (short-term)	72 per 1000	<b>8 per 1000</b> (1 to 43)	<b>RR 0.11</b> (0.02 to 0.59)	360 (2 studies)	⊕⊕⊝⊝ low <sup>1,2</sup>	Relapse was not reported but 'no clini- cally important change in global state' was reported and the other outcome with different criteria of 'no clinically important change in global state' indi- cated no difference between the two comparison groups in the short-term.
Mental state: No clinically im- portant change in mental state	See comment	See comment	Not estimable	-	See comment	Not reported.
Behaviour: Leaving the study early (short-term)	See comment	See comment	Not estimable	454 (3 studies)	⊕⊕⊝© low <sup>1,2</sup>	No participants left either group early.
Service outcomes: Hospitalisa- tion	See comment	See comment	Not estimable	-	See comment	Not reported.
Adverse effects: Clinically im- portant general adverse effects	See comment	See comment	Not estimable	0 (0)	See comment	Three studies reported this outcome, however, two studies only reported one group's data and the other one study did not report the data.
Quality of life: No clinically im- portant change in quality of life	See comment	See comment	Not estimable	-	See comment	Not reported.
Economic outcomes: Cost of care	See comment	See comment	Not estimable	-	See comment	Not reported.
*The basis for the <b>assumed risk</b> (e. based on the assumed risk in the co <b>CI:</b> Confidence interval; <b>RR:</b> Risk ra	.g. the median cont omparison group a tio;	rol group risk across nd the <b>relative effe</b>	s studies) is provide <b>ct</b> of the interventi	ed in footnotes. The on (and its 95% CI).	e corresponding ris	<b>k</b> (and its 95% confidence interval) is

GRADE Working Group grades of evidence

High quality: Further research is very unlikely to change our confidence in the estimate of effect.

**Moderate quality:** Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate. **Low quality:** Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate. **Very low quality:** We are very uncertain about the estimate.

œ

Cochrane

Trusted evidence. Informed decisions. Better health.

# Summary of findings 5. ACUPUNCTURE versus TCM DRUG for schizophrenia

# ACUPUNCTURE versus TCM DRUG for schizophrenia

**Patient or population:** patients with schizophrenia **Intervention:** ACUPUNCTURE versus TCM DRUG

Outcomes	Illustrative comparative risks* (95% CI)		Relative effect (95% CI)	No of Partici- pants (studies)	Quality of the evidence (GRADF)	Comments	
	Assumed risk	Corresponding risk		(studies)	(0.0.0_)		
	Control	ACUPUNCTURE versus TCM DRUG					
Global state: Not improved, endpoint (short-term) - Not im- proved (no change in symp- toms) - Electroacupuncture	711 per 1000	<b>370 per 1000</b> (242 to 569)	<b>RR 0.52</b> (0.34 to 0.80)	88 (1 study)	⊕⊕⊝⊝ low <sup>1,2</sup>	Relapse was not reported but 'no clini- cally important change in global state' was reported and other outcomes with different criteria of 'no clinically im- portant change in global state' indi- cated no difference between the two comparison groups in the short-term.	
Mental state: No clinically im- portant change in general men- tal state	See comment	See comment	Not estimable	-	See comment	Not reported.	
Behaviour: Leaving the study early (short-term)	See comment	See comment	Not estimable	328 (3 studies)	See comment	No participants left either group early.	
Service outcomes: Hospitalisa- tion	See comment	See comment	Not estimable	-	See comment	Not reported.	
Adverse effects: Clinically im- portant general adverse effects	See comment	See comment	Not estimable	-	See comment	Three studies reported this outcome, however, two studies only reported one group's data and the other one study did not report the data.	
Quality of life: No clinically im- portant change in quality of life	See comment	See comment	Not estimable	-	See comment	Not reported.	

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

<b>Cl:</b> Confidence interval; <b>RR:</b> Risk ratio;							
GRADE Working Group grades of evidenc High quality: Further research is very ur Moderate quality: Further research is lil Low quality: Further research is very lik Very low quality: We are very uncertain	e nlikely to change ou kely to have an imp ely to have an impo about the estimate	ur confidence in the estimate portant impact on our confide prtant impact on our confide e.	e of effect. lence in the estimate ence in the estimate	e of effect and may of effect and is like	v change the estimat ely to change the est	e. imate.	
Risk of bias: rated 'serious' Unblinding mpression: rated 'serious' - Few particip ummary of findings 6. ELECTRIC A	of participants and pants and/or few ev	d personnel. ens. <b>CONVULSIVE THERAPY ve</b>	ersus ELECTROCO	NVULSIVE THEF	RAPY for schizoph	renia	
LECTRIC ACUPUNCTURE CONVULSIVE	E THERAPY versus	ELECTRIC CONVULSIVE TH	ERAPY for schizopl	nrenia			
Patient or population: patients with scl Intervention: ELECTRIC ACUPUNCTURE	hizophrenia CONVULSIVE THE	RAPY versus ELECTRIC CONV	ULSIVE THERAPY				
Outcomes	Illustrative comparative risks* (95% CI)		Relative effect	No of Partici-	Quality of the	Comments	
			(95% CI)	nante	ovidonco		
	Assumed risk	Corresponding risk	– (95% CI)	pants (studies)	evidence (GRADE)		
	Assumed risk Control	Corresponding risk ELECTRIC ACUPUNC- TURE CONVULSIVE THERAPY versus ELEC- TRIC CONVULSIVE THERAPY	– (95% CI) –	pants (studies)	evidence (GRADE)		
Global state: Relapse	Assumed risk Control See comment	Corresponding risk ELECTRIC ACUPUNC- TURE CONVULSIVE THERAPY versus ELEC- TRIC CONVULSIVE THERAPY See comment	(95% CI) Not estimable	pants (studies) -	evidence (GRADE) See comment	Not reported.	
Global state: Relapse Mental state: No clinically important change in general mental state	Assumed risk Control See comment See comment	Corresponding risk ELECTRIC ACUPUNC- TURE CONVULSIVE THERAPY versus ELEC- TRIC CONVULSIVE THERAPY See comment See comment	(95% CI) Not estimable Not estimable	pants (studies) - -	evidence (GRADE) See comment See comment	Not reported. Not reported.	
Global state: Relapse Mental state: No clinically important change in general mental state Behaviour: Leaving the study early [short-term]	Assumed risk Control See comment See comment See comment	Corresponding risk ELECTRIC ACUPUNC- TURE CONVULSIVE THERAPY versus ELEC- TRIC CONVULSIVE THERAPY See comment See comment See comment See comment	<ul> <li>(95% CI)</li> <li>Not estimable</li> <li>Not estimable</li> <li>Not estimable</li> </ul>	pants (studies)	evidence (GRADE) See comment See comment ⊕⊕⊙⊙ low 1,2	Not reported. Not reported. Not reported. No participants in either group left early.	

Economic outcomes: Cost of

care

See comment

See comment

Not estimable

-

See comment

Not reported.

# Cochrane Library

Trusted evidence. Informed decisions. Better health.

Adverse effects: Spinal fracture (short-term)	441 per 1000	<b>146 per 1000</b> (62 to 357)	<b>RR 0.33</b> (0.14 to 0.81)	68 (1 study)	⊕⊕⊙© low <sup>1,2</sup>	Data from a single study where there was no dif- ference in back pain be- tween the two groups.	
Quality of life: No clinically impor- tant change in quality of life	See comment	See comment	Not estimable	-	See comment	Not reported.	
Economic outcomes: Cost of care	See comment	See comment	Not estimable	-	See comment	Not reported.	
*The basis for the <b>assumed risk</b> (e.g. the median control group risk across studies) is provided in footnotes. The <b>corresponding risk</b> (and its 95% confidence interval) is based on the assumed risk in the comparison group and the <b>relative effect</b> of the intervention (and its 95% CI).							
GRADE Working Group grades of evidence High quality: Further research is very unlikely to change our confidence in the estimate of effect. Moderate quality: Further research is likely to have an important impact on our confidence in the estimate of effect and may change the estimate. Low quality: Further research is very likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate. Very low quality: We are very uncertain about the estimate.							

<sup>1</sup> Risk of bias; rated 'serious' - Unblinding of participants and personnel. <sup>2</sup> Imprecision: rated 'serious' - Few participants and few events. 4.4440-

Cochrane Library

Trusted evidence. Informed decisions. Better health.



# BACKGROUND

# **Description of the condition**

Schizophrenia affects 1% of people and therefore, one fifth of people with schizophrenia are of Chinese origin. In Chinese health care, theories for the aetiology, pathology and treatment of schizophrenia are different to those of Western medicine. Conventional medicine diagnoses schizophrenia primarily by operationalised criteria such as the Diagnostic and Statistical Manual (DSM-IV) or the International Classification of Diseases (ICD10). Chinese medicine uses different methodology to diagnose mental health disorders such as schizophrenia by pattern differentiation. The four diagnostic methods (inspection, listening/ smelling, inquiry and palpation) are the way by which patient information is obtained (Xu 1991). This information is then analysed using one or several diagnostic models (Zang Fu theory, Eight Principles [yin/yang, interior/exterior, excess/deficiency, hot/cold], Four Division Pattern [Wei Qi Ying Xue ], Six Division Pattern, Five Elements Pattern [wood, fire, earth, metal, and water], Pattern of Qi, Blood, and Body Fluids, Three Burners Pattern) to arrive at a diagnosis. Thus, two people diagnosed with DSM-IV schizophrenia could nevertheless, from a Chinese medicine perspective, have different sub-types or patterns and therefore require different treatment. The Psychosis Professional Committee of Chinese Integrative Medicine Association drafted the Standard of Integrative Medicine Syndrome Type of Schizophrenia. There

### Figure 1. Acupuncture

are six main patterns which fall within the disease category of Dian Kuang/withdrawal mania which can also encompass schizophrenia. The six types are: 1. Internal disturbance of pyrophlegm; 2. Internal retention of phlegm and dampness; 3. Qi stagnation and blood stasis; 4. Yin deficiency and fire excess; 5. Yang deficiency; 6.Other miscellaneous types (The Psychosis Professional Committee 1988; Guo 2010; Zhang 1996).

# **Description of the intervention**

The illness often causes distortions of perception, thinking, behaving and even movement and, although antipsychotic drugs have been the mainstay of treatment of schizophrenia since the early 1950s, these treatments still leave many with residual symptoms and disabling adverse effects. Acupuncture (Figure 1) has been shown to have few adverse effects (Ernst 2001; MacPherson 2001) and may be more socially acceptable, tolerable and inexpensive than the more conventional drugs available from the pharmaceutical industry. Chinese medicine, which includes acupuncture and is also referred to as Traditional Chinese Medicine (TCM), has been used to treat 'schizophrenia-like illnesses' (i.e. Dian Kuang/withdrawal mania) for over 2000 years (Ming 2001). Acupuncture is practiced as an accepted healthcare model in China, Korea and Japan, although the methods used in each country are distinct (Kaptchuk 2002). There are many categories of acupuncture such as traditional acupuncture, electroacupuncture, laser acupuncture, and acupoint injection.



# How the intervention might work

Acupuncture involves the stimulation of specific points (acupoints). The internal diseases are treated using external applications by dredging meridians, regulating Yin and Yang, reinforcing the healthy Qi and eliminating the pathogenic factors. Though acupuncture has a long history, any mechanism for treating schizophrenia is still unclear (Shi 2010; Xu 2010). Acupuncture may affect the central nervous functions of the cerebral cortex through recuperating the Qi, blood, dredging meridians and then adjusting the nervous system's functions and endocrine level (Wang 2007). There is, however, not enough modern neural physiological and biochemical research, imaging or animal studies to support this (Xu 2010).

#### Why it is important to do this review

Acupuncture has been used to treat schizophrenia for many years and now it is considered as one of the most popular types of complementary and alternative medicine (CAM) in Western health care. We wish to update past work, undertake a review that is systematic and quantify the effects of this inexpensive treatment.

# OBJECTIVES

To review the effects of acupuncture for people with schizophrenia and related psychoses, evaluating acupuncture alone or in combination regimens compared with placebo (or no treatment), or any other treatments.



# METHODS

# Criteria for considering studies for this review

# **Types of studies**

We included all relevant randomised controlled trials. We planned that if a trial was described as 'double-blind' but it was implied that the study was randomised, these trials would have been included in a sensitivity analysis. If there was no substantive difference within primary outcomes (see 'Types of outcome measures') when these 'implied randomisation' studies were added, then they would have been included in the final analysis. If there was a substantive difference, only clearly randomised trials would have been utilised and the results of the sensitivity analysis described in the text. Quasi-randomised studies, such as those allocating by using alternate days of the week, were excluded.

# **Types of participants**

We included patients with schizophrenia, schizophreniform psychosis and schizophrenia-like illnesses, any age, diagnosed by any criteria.

# **Types of interventions**

# 1. All categories of acupuncture

Administered solely or in conjunction with any other treatments, versus:

# 2. Placebo (sham acupuncture) or no treatment

# 3. Any other treatments

We do recognise that there are other possible combinations of treatments that could be included in this review but this (2012) update focused on only the above comparisons.

For previous type of interventions please see Appendix 1.

#### Types of outcome measures

Outcomes were divided into short-term (less than three months), medium-term (three to 12 months) and long-term (more than one year).

#### **Primary outcomes**

# 1. Global state

1.1 Relapse

# 2. Mental state

2.2 No clinically important change in general mental state - as defined by each of the studies

# 3. Service outcomes

3.1 Hospitalisation

# 4. Adverse effects

4.1 Clinically important general adverse effects

# Secondary outcomes

# 1. Death - suicide or natural causes

# 2. Global state

2.1 No clinically important change in global state - as defined by each of the studies

- 2.2 Average endpoint global state score
- 2.3 Average change in global state scores

# 3. Mental state

- 3.1 No change in general mental state
- 3.2 Average endpoint general mental state score
- 3.3 Average change in general mental state scores
- 3.4 No clinically important change in specific symptoms as
- defined by each of the studies
- 3.5 No change in specific symptoms
- 3.6 Average endpoint specific symptom score
- 3.7 Average change in specific symptom scores

# 4. Behaviour

4.1 Leaving the study early

4.2 No clinically important change in general behaviour - as defined by each of the studies

- 4.3 No change in general behaviour
- 4.4 Average endpoint general behaviour score
- 4.5 Average change in general behaviour scores

4.6 No clinically important change in specific aspects of behaviour

- 4.7 No change in specific aspects of behaviour
- 4.8 Average endpoint specific aspects of behaviour
- 4.9 Average change in specific aspects of behaviour

# 5. Service outcomes

5.1 Time to hospitalisation

# 6. Adverse effects

- 6.1 Any general adverse effects
- 6.2 Average endpoint general adverse effect score
- 6.3 Average change in general adverse effect scores

6.4 No clinically important change in specific adverse effects - as

defined by each of the studies

6.5 No change in specific adverse effects

- 6.6 Average endpoint specific adverse effects
- 6.7 Average change in specific adverse effects

# 7. Engagement with services

7.1 No clinically important engagement - as defined by each of the studies

- 7.2 No engagement
- 7.3 Average endpoint engagement score
- 7.4 Average change in engagement scores

# 8. Satisfaction with treatment

- 8.1 Recipient of care not satisfied with treatment
- 8.2 Recipient of care average satisfaction score
- 8.3 Recipient of care average change in satisfaction scores

# 9. Quality of life

9.1 No clinically important change in quality of life - as defined by each of the studies9.2 No change in quality of life

Acupuncture for schizophrenia (Review)

Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



9.3 Average endpoint quality of life score9.4 Average change in quality of life scores

#### 10. Economic outcomes

10.1 Costs of care

# 11. 'Summary of findings' table

We used the GRADE approach to interpret findings (Schünemann 2008) and used the GRADE profiler to import data from Review Manager (RevMan) to create 'Summary of findings' tables. These tables provide outcome-specific information concerning the overall quality of evidence from each included study in the comparison, the magnitude of effect of the interventions examined, and the sum of available data on all outcomes we rated as important to patient-care and decision making. We selected the following main outcomes for inclusion in the 'Summary of findings' table.

1. Global state: relapse.

2. Mental state: no clinically important change in general mental state - as defined by each of the studies.

3. Behaviour: leaving the study early.

4. Service outcomes: hospitalisation.

5. Adverse effects: clinically important general adverse effects.

6. Quality of life: no clinically important change in quality of life - as defined by each of the studies.

7. Economic outcomes: costs of care.

# Search methods for identification of studies

# **Electronic searches**

# 1. Cochrane Schizophrenia Group's Trials Register

The Trials Search Co-ordinator (TSC) searched the Cochrane Schizophrenia Group's Registry of Trials (February 2012) using the following search strategies:

(\*acup\* or \*moxibustion\*) in Title Field of REFERENCE or (\*acupuncture\* or \*moxibustion\*) in Intervention Field of STUDY

The Cochrane Schizophrenia Group's Registry of Trials is compiled by systematic searches of major resources (including AMED, BIOSIS, CINAHL, EMBASE, MEDLINE, PsycINFO, PubMed, and registries of clinical trials) and their monthly updates, handsearches, grey literature, and conference proceedings (see Group Module). There is no language, date, document type, or publication status limitations for inclusion of records into the register.

For previous searches, see Appendix 2.

#### Searching other resources

# 1. Reference searching

We inspected references of all identified studies for further relevant studies.

# 2. Personal contact

We contacted the first author of each included study for information regarding unpublished trials.

# Data collection and analysis

Methods used in data collection and analysis for this 2012 update are below; for previous methods, please see Appendix 3.

# Selection of studies

For this 2012 update, review author XS independently inspected citations from the new electronic search and identified relevant abstracts. XS also inspected full articles of the abstracts meeting inclusion criteria. JX carried out the reliability check of all citations from the new electronic search. Where disputes arose, we resolved disagreements by discussion or, if doubt remained, by acquiring the full article for more detailed scrutiny. If doubt still remained, we added these trials to the list of those awaiting assessment pending acquisition of further information and we contacted the authors of studies. For details of previous author contributions in study selection see Acknowledgements.

# Data extraction and management

# 1. Extraction

For this 2012 update, XS extracted data from all included studies. In addition, to ensure reliability, JX independently extracted data from a random sample of these studies, comprising 10% of the total. We discussed disagreements, documented our decisions and, if necessary, contracted authors of studies for clarification. With any remaining problems, CEA helped clarify issues, although he could not translate the Mandarin text. When uncertainty persisted, we allocated the trial to Studies awaiting classification. Data presented only in graphs and figures were extracted whenever possible, but included only if two review authors independently extracted the same results. In order to obtain missing information or for clarification, whenever necessary we contacted authors through an open-ended request . If studies were multi-centre, where possible, we extracted data relevant to each component centre separately.

# 2. Management

# 2.1 Forms

We extracted data onto standard, simple forms.

#### 2.2 Scale-derived data

We included continuous data from rating scales only if: a. the psychometric properties of the measuring instrument have been described in a peer-reviewed journal (Marshall 2000); and b. the measuring instrument has not been written or modified by one of the trialists for that particular trial.

Ideally, the measuring instrument should either be i. a self-report or ii. completed by an independent rater or relative (not the therapist). We realise that this is not often reported clearly; we have noted whether or not this is the case in Description of studies.

#### 2.3 Endpoint versus change data

There are advantages of both endpoint and change data. Change data can remove a component of between-person variability from the analysis. On the other hand, calculation of change needs two assessments (baseline and endpoint), which can be difficult in unstable and difficult to measure conditions such as schizophrenia. We decided primarily to use endpoint data, and only use change data if the former were not available. We combined endpoint and change data in the analysis as we used mean differences (MD)

Acupuncture for schizophrenia (Review)

Copyright  ${\small ©}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



rather than standardised mean differences throughout (Higgins 2011, Chapter 9.4.5.2).

Previous version of this review did not combine endpoint and change data in the analysis (see Appendix 3).

# 2.4 Skewed data

Continuous data on clinical and social outcomes are often not normally distributed. To avoid the pitfall of applying parametric tests to non-parametric data, we aimed to apply the following standards to all data before inclusion:

a) standard deviations (SDs) and means are reported in the paper or obtainable from the authors;

b) when a scale starts from the finite number zero, the SD, when multiplied by two, is less than the mean (as otherwise the mean is unlikely to be an appropriate measure of the centre of the distribution, (Altman 1996);

c) if a scale started from a positive value (such as the Positive and Negative Syndrome Scale (PANSS), (Kay 1986)), which can have values from 30 to 210), the calculation described above will be modified to take the scale starting point into account. In these cases skew is present if 2 SD > (S-S min), where S is the mean score and 'S min' is the minimum score.

Endpoint scores on scales often have a finite start and end point and these rules can be applied. Skewed data pose less of a problem when looking at means if the sample size is large (>200) and we will enter these into the syntheses. We presented skewed endpoint data from studies of less than 200 participants as 'Other data' in the Data & analyses rather than enter such data in analyses.

When continuous data are presented on a scale that includes a possibility of negative values (such as change data), it is difficult to tell whether data are skewed or not. We planned to present and enter change data into analyses.

#### 2.5 Common measure

To facilitate comparison between trials, we intended to convert variables that can be reported in different metrics, such as days in hospital (mean days per year, per week or per month) to a common metric (e.g. mean days per month).

#### 2.6 Conversion of continuous to binary

Where possible, we made efforts to convert outcome measures to dichotomous data. This can be done by identifying cut-off points on rating scales and dividing participants accordingly into 'clinically improved' or 'not clinically improved'. It is generally assumed that if there is a 50% reduction in a scale-derived score such as the Brief Psychiatric Rating Scale (BPRS, Overall 1962) or the Positive and Negative Syndrome Scale (PANSS, Kay 1986), this could be considered as a clinically significant response (Leucht 2005; Leucht 2005a). If data based on these thresholds were not available, we used the primary cut-off presented by the original authors.

#### 2.7 Direction of graphs

Where possible, we entered data in such a way that the area to the left of the line of no effect indicated a favourable outcome for acupuncture. If we had to enter data so the area to the left of the line indicated a favourable outcome for the control treatment, this was noted in the relevant graphs.

#### Assessment of risk of bias in included studies

For this 2012 update, XS worked independently by using criteria described in the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011) to assess trial quality. This new set of criteria is based on evidence of associations between overestimate of effect and high risk of bias of the article such as sequence generation, allocation concealment, blinding, incomplete outcome data and selective reporting. JX assessed risk of bias from a random sample of these studies, comprising 10% of the total.

Any disagreements between raters were resolved through discussion with CEA. Where inadequate details of randomisation and other characteristics of trials were provided, we contacted authors of the studies in order to obtain further information. All disagreements in quality assessment were recorded and reported. If disputes arose as to which quality category a trial was to be rated, again, we resolved these through discussion.

We have noted the level of risk of bias in both the text of the review and in the 'Summary of findings' tables.

The previous version of this review used a different, less welldeveloped, means of categorising risk of bias (see Appendix 3).

#### Measures of treatment effect

# 1. Binary data

For binary outcomes we calculated a standard estimation of the risk ratio (RR) and its 95% confidence interval (CI). It has been shown that RR is more intuitive (Boissel 1999) than odds ratios and that odds ratios tend to be interpreted as RR by clinicians (Deeks 2000). The Number Needed to Treat/Harm (NNT/H) statistic with its confidence intervals is intuitively attractive to clinicians but is problematic both in its accurate calculation in meta-analyses and interpretation (Hutton 2009). For binary data presented in the 'Summary of findings' tables, where possible, we calculated illustrative comparative risks.

# 2. Continuous data

For continuous outcomes we estimated mean difference (MD) between groups. We would prefer not to calculate effect size measures (standardised mean difference (SMD)). However, if scales of very considerable similarity were used, we presumed there was a small difference in measurement, and we would have calculated effect size and transformed the effect back to the units of one or more of the specific instruments.

#### Unit of analysis issues

# 1. Cluster trials

Studies increasingly employ 'cluster randomisation' (such as randomisation by clinician or practice), but analysis and pooling of clustered data poses problems. Authors often fail to account for intra-class correlation in clustered studies, leading to a 'unit of analysis' error (Divine 1992) whereby P values are spuriously low, confidence intervals unduly narrow and statistical significance overestimated. This causes type I errors (Bland 1997; Gulliford 1999).

If cluster trials where clustering was not accounted for in primary studies had been included, we would have presented data in a table, with a (\*) symbol to indicate the presence of a probable

Acupuncture for schizophrenia (Review)

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Cochrane Library

Trusted evidence. Informed decisions. Better health.

Cochrane Database of Systematic Reviews

unit of analysis error. We would have contacted the first authors of studies to obtain intra-class correlation coefficients (ICCs) for their clustered data and to adjust for this by using accepted methods (Gulliford 1999). If clustering had been incorporated into the analysis of primary studies, we would have presented these data as if from a non-cluster randomised study, but adjusted for the clustering effect.

We have sought statistical advice and have been advised that the binary data as presented in a report should be divided by a 'design effect'. This is calculated using the mean number of participants per cluster (m) and the ICC [Design effect = 1+(m-1)\*ICC] (Donner 2002). If the ICC had not been reported, it would have been assumed to be 0.1 (Ukoumunne 1999).

If cluster studies have been appropriately analysed taking into account ICCs and relevant data documented in the report, synthesis with other studies would have been possible using the generic inverse variance technique.

# 2. Cross-over trials

A major concern of cross-over trials is the carry-over effect. It occurs if an effect (e.g. pharmacological, physiological or psychological) of the treatment in the first phase is carried over to the second phase. As a consequence, on entry to the second phase, the participants can differ systematically from their initial state despite a washout phase. For the same reason cross-over trials are not appropriate if the condition of interest is unstable (Elbourne 2002). As both effects are very likely in severe mental illness, if we had identified any cross-over trials, we would only have used data from the first phase of cross-over studies.

# 3. Studies with multiple treatment groups

Where a study involves more than two treatment arms, if relevant, we would have presented the additional treatment arms in comparisons. If data were binary we would simply add these and combine within the two-by-two table. If data were continuous we would have combined data following the formula in section 7.7.3.8 (Combining groups) of the *Handbook* (Higgins 2011). Where the additional treatment arms were not relevant, we did not reproduce these data.

# Dealing with missing data

# 1. Overall loss of credibility

At some degree of loss of follow-up, data must lose credibility (Xia 2009). We chose that, if for any particular outcome, should more than 50% of data be unaccounted for, we would not reproduce these data or use them within analyses, (except for the outcome 'leaving the study early'). If, however, more than 50% of those in one arm of a study were lost, but the total loss was less than 50%, we would mark such data with (\*) to indicate that such a result may well be prone to bias.

The previous version of this review excluded data from studies where more than 50% of participants in any group were lost to follow-up (see Appendix 3).

# 2. Binary

In the case where attrition for a binary outcome was between 0% and 50% and where these data were not clearly described, we presented data on a 'once-randomised-always-analyse' basis (an

intention-to-treat analysis). Those leaving the study early were all assumed to have the same rates of negative outcome as those who completed, with the exception of the outcome of death and adverse effects. For these outcomes, the rate of those who stayed in the study - in that particular arm of the trial - was used for those who did not. If the above assumptions had been used for the primary outcomes we would have undertaken a sensitivity analysis to test how prone the primary outcomes were to change when data only from people who completed the study to that point were compared to the intention-to-treat analysis using the above assumptions.

# 3. Continuous

### 3.1 Attrition

In the case where attrition for a continuous outcome was between 0% and 50%, and data only from people who completed the study to that point were reported, we reproduced these.

#### 3.2 Standard deviations

When standard deviations were not reported, we first tried to obtain the missing values from the authors. When not available, where there are missing measures of variance for continuous data, but an exact standard error (SE) and confidence intervals are available for group means, and either a P value or T value available for differences in mean, we could calculate them according to the rules described in the Handbook (Higgins 2011): When only the SE is reported, standard deviations (SDs) are calculated by the formula SD = SE \* square root (n). Chapters 7.7.3 and 16.1.3 of the Handbook (Higgins 2011) present detailed formulae for estimating SDs from P values, T or F values, confidence intervals, ranges or other statistics. If these formulae do not apply, we can calculate the SDs according to a validated imputation method which is based on the SDs of the other included studies (Furukawa 2006). Although some of these imputation strategies can introduce error, the alternative would be to exclude a given study's outcome and thus to lose information. If possible, we planned to examine the validity of the imputations in a sensitivity analysis excluding imputed values.

#### 3.3 Last observation carried forward

We anticipated that in some studies the method of last observation carried forward (LOCF) would be employed within the study report. As with all methods of imputation to deal with missing data, LOCF introduces uncertainty about the reliability of the results (Leucht 2007). Therefore, if LOCF data had been used in the trial, if less than 50% of the data had been assumed, we would have reproduced these data and indicated that they were the product of LOCF assumptions.

#### Assessment of heterogeneity

# 1. Clinical heterogeneity

We considered all included studies initially, without seeing comparison data, to judge clinical heterogeneity. We simply inspected all studies for clearly outlying people or situations which we had not predicted would arise. When such situations or participant groups arose, we fully discussed these.

# 2. Methodological heterogeneity

We considered all included studies initially, without seeing comparison data, to judge methodological heterogeneity. We simply inspected all studies for clearly outlying methods which we

Acupuncture for schizophrenia (Review)

Copyright  $\odot$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

had not predicted would arise. When such methodological outliers arose, we fully discussed these.

#### 3. Statistical heterogeneity

#### 3.1 Visual inspection

We visually inspected graphs to investigate the possibility of statistical heterogeneity.

#### 3.2 Employing the I<sup>2</sup> statistic

We investigated heterogeneity between studies by considering the I<sup>2</sup> method alongside the Chi<sup>2</sup> P value. The I<sup>2</sup> provides an estimate of the percentage of inconsistency thought to be due to chance (Higgins 2003). The importance of the observed value of I<sup>2</sup> depends on i. magnitude and direction of effects and ii. strength of evidence for heterogeneity (e.g. P value from Chi<sup>2</sup> test, or a confidence interval for I<sup>2</sup>). An I<sup>2</sup> estimate greater than or equal to around 50% accompanied by a statistically significant Chi<sup>2</sup> statistic was interpreted as evidence of substantial levels of heterogeneity (Higgins 2011). When substantial levels of heterogeneity were found in the primary outcome, we explored reasons for heterogeneity (Subgroup analysis and investigation of heterogeneity).

The previous version of this review used a different, less-well developed approach to deal with heterogeneity (see Appendix 3).

# **Assessment of reporting biases**

Reporting biases arise when the dissemination of research findings is influenced by the nature and direction of results (Egger 1997). These are described in Section 10 of the *Handbook* (Higgins 2011). We are aware that funnel plots may be useful in investigating reporting biases but are of limited power to detect small-study effects. We did not use funnel plots for outcomes where there were 10 or fewer studies, or where all studies were of similar sizes. In other cases, where funnel plots are possible, we sought statistical advice in their interpretation.

The previous version of this review used a different, less-well developed approach to assess the reporting biases (see Appendix 3).

#### **Data synthesis**

We understand that there is no closed argument for preference for use of fixed-effect or random-effects models. The random-effects method incorporates an assumption that the different studies are estimating different, yet related, intervention effects. This often seems to be true to us and the random-effects model takes into account differences between studies even if there is no statistically significant heterogeneity. There is, however, a disadvantage to the random-effects model: it puts added weight onto small studies, which often are the most biased ones. Depending on the direction of effect, these studies can either inflate or deflate the effect size. We chose the fixed-effect model for all analyses. we only used random-effects model when heterogeneity was present.

#### Subgroup analysis and investigation of heterogeneity

#### 1. Subgroup analyses

#### 1.1 Clinical state, stage or problem

We proposed to undertake this review and provide an overview of the effects of acupuncture for people with schizophrenia in general. In addition, however, if possible, we tried to report data on subgroups of people in the same clinical state, stage and with similar problems.

#### 2. Investigation of heterogeneity

If inconsistency was high, we have reported this. First, we investigated whether data had been entered correctly. Second, if data were correct, we visually inspected the graph and successively removed outlying studies to see if homogeneity was restored. For this review, we decided that should this occur with data contributing to the summary finding of no more than around 10% of the total weighting, we would present data. If not, then we did not pool data and discussed issues. We know of no supporting research for this 10% cut-off, but we use prediction intervals as an alternative to this unsatisfactory state.

When unanticipated clinical or methodological heterogeneity was obvious, we simply stated hypotheses regarding these for future reviews or versions of this review. We do not anticipate undertaking analyses relating to these.

# Sensitivity analysis

#### 1. Implication of randomisation

Had we found trials which were described in some way so as to imply randomisation, we would have included these in a sensitivity analysis. For the primary outcomes we would have included these studies and if there was no substantive difference when the implied randomised studies were added to those with better description of randomisation, then we would have entered all data from these studies.

# 2. Assumptions for lost binary data

Where assumptions had to be made regarding people lost to followup (see Dealing with missing data), we compared the findings of the primary outcomes when we used our assumption/s and when we used data only from people who completed the study to that point. If there was a substantial difference, we reported results and discussed them, but continued to employ our assumption.

Where assumptions had to be made regarding missing SDs data (see Dealing with missing data), if possible, we compared the findings of outcomes when we used our assumption/s and when we used data only from people who completed the study to that point. If there was a substantial difference, we reported results and discussed them, but continued to employ our assumption.

#### 3. Risk of bias

We analysed the effects of excluding trials that were judged to be at high risk of bias across one or more of the domains of randomisation (implied as randomised with no further details available): allocation concealment, blinding and outcome reporting for the meta-analysis of the primary outcome. If the exclusion of trials at high risk of bias did not substantially alter the direction of effect or the precision of the effect estimates, then we included data from these trials in the analysis.

# 4. Imputed values

If we had found cluster randomised trials we would have undertaken a sensitivity analysis to assess the effects of including data from trials where we used imputed values for ICC in calculating the design effect in cluster randomised trials.

If we had undertaken sensitivity analysis and noted substantial differences in the direction or precision of effect estimates in any of the sensitivity analyses listed above, we would not have pooled data from the excluded trials with the other trials contributing to the outcome, but would have presented them separately.

# RESULTS

# **Description of studies**

For substantive descriptions of studies please see Characteristics of included studies, Characteristics of excluded studies and Characteristics of studies awaiting classification.

Previous version of description of studies please see Appendix 4.

# **Results of the search**

The update searches in 2012 yielded references to 49 reports including all the reports in the previous version (Figure 2). We selected 39 studies with 47 reports for further inspection of full papers excluding 2 records for one not relating to acupuncture and one not relating with schizophrenia after initial appraisal. We were able to include 30 studies with 36 reports.



# Figure 2. Study flow diagram.



#### **Included studies**

We included thirty studies.

# 1. Allocation

All included studies were randomised controlled trials. Only 10 of the included studies described the randomisation method. Three studies used a random sampling method (acupoint cat - Sun 2005; electro - Chen 2008; electro - Zhang 2001); three were

randomised by using a random allocation table (traditional - Liu 2010; traditional - Wang 2006; traditional - Xu 2004); two drew lots (acupoint inj - Wang 2000; electro - Yao 2006); one study used the SAS program - this was the only trial that described allocation concealment by using opaque sealed envelopes (electro - Cheng 2009) and one used the coin-tossing method (electro - Cui 2000). Five studies were assessor-blinded (acupoint inj - Pan 2002; acupoint inj - Wang 2000; electro - Xiong 2010; electro - Zhou 1997; laser - Zhang 1991); two were patient- and assessor-blinded (electro

Acupuncture for schizophrenia (Review)

Copyright  ${\small ©}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Cochrane Library

Trusted evidence. Informed decisions. Better health.

- Cheng 2009; traditional - Bouhlel 2011); and we could not ascertain how blinding was achieved in traditional - Liu 2010, although it was reported as single blind.

# 2. Study duration

Study duration varied from two weeks (acupoint inj - Wang 2000) to six months (acupoint inj - Yang 2000). The majority of studies were short-term. Five of all included studies were medium-term (acupoint inj - Yang 2000; electro - Chen 2008; traditional - Liu 2010; traditional - Luo 2006; traditional - Tang 2005) and the duration of laser - Ma 1999 was unclear. acupoint inj - Pan 2002 had a two-year follow-up and electro - Yao 2006 a six-month follow-up.

# 3. Setting

Twenty-two studies were hospital-based with inpatients. Two studies, electro - Chen 2006 and traditional - Ma 2008, were undertaken with both inpatients and outpatients. Two studies were undertaken with outpatients (traditional - Zhao 2005a; traditional - Zhao 2005b). The settings of the four remaining studies were not mentioned (acupoint inj - Yang 2000; laser - Liu 1986; traditional - Luo 2006; EACT - Xue 1987).

#### 4. Country

All included studies were undertaken in China except one traditional - Bouhlel 2011, which took place in Tunisia.

#### 5. Participants

It was reported in all studies that the participants suffered from schizophrenia. Seventeen adopted the standards of Chinese Classification of Mental Disorder such as second edition (CCMD-2, acupoint inj - Yang 2000; laser - Ma 1999), second edition revision (CCMD-2-R, acupoint inj - Pan 2002; acupoint cat - Wang 1997; acupoint inj - Wang 2000; traditional - Wang 2006; electro - Cui 2000; electro - Zhang 1993; traditional - Xu 2004) or third edition (CCMD-3, acupoint cat - Sun 2005; electro - Chen 2006; electro -Ding 2005; electro - Xiong 2010; electro - Yao 2006; traditional - Liu 2010; traditional - Luo 2006; traditional - Tang 2005). Two studies diagnosed schizophrenia according to DSM IV (electro - Cheng 2009; traditional - Bouhlel 2011). Two studies diagnosed schizophrenia with both CCMD-3 and Andreasen's diagnosis standards (electro - Chen 2008; electro - Wang 2005); two with both standards of CCMD-2-R and DSM III (laser - Zhang 1991) or CCMD-2-R and ICD-10 (electro - Zhang 2001); three with both standards of TCM and CCMD-2-R (traditional - Zhao 2005a; traditional - Zhao 2005b) or TCM and CCMD-3 (traditional - Ma 2008). Only one study (electro -Zhou 1997) adopted three standards - DSM III, CCMD and TCM. The other three studies did not mention the diagnostic standard (EACT - Xue 1987; electro - Zhang 1987; laser - Liu 1986).

The age of participants of 15 studies ranged from 15 to 67 years. Three studies did not definitely describe the age range (traditional - Tang 2005; traditional - Zhao 2005a; traditional - Zhao 2005b). The other 12 studies reported mean age. The majority of studies included both women and men except electro - Ding 2005 and traditional - Xu 2004 (only men). acupoint inj - Pan 2002 and electro - Zhang 1993 did not describe gender. All studies except traditional - Tang 2005 reported the history of illness.

# 6. Study size

The number of participants ranged from 31 to 300; acupoint cat -Wang 1997, traditional - Zhao 2005a and traditional - Zhao 2005b were over 200 participants. The size of electro - Zhang 1993 was unclear.

# 7. Interventions

We found six categories of acupuncture were used whether alone or in combination regimens: traditional acupuncture, electroacupuncture, acupoint injection, laser acupuncture, acupoint catgut treatment and electric acupuncture convulsive therapy. Traditional acupuncture included acupuncture manipulation and moxibustion. Acupuncture manipulation refers to an operation method to prevent disease by using different needles or non-needle way to stimulate the specific points (acupoints) with certain practices or methods. Moxibustion not only refers to burning, smoking and ironing body surface mainly using moxa but also refers to any external treatment of non-fire source (Shi 2007). Electroacupuncture refers to a combination method of needles and electrical stimulation to prevent disease that connects needles with trace current which are close to the human bioelectricity after inserting needles into acupoints and getting the feeling of 'DeQi' (Shi 2007). Acupoint injection, guiding by the basic theory of Chinese medicine, is a treatment method with synergistic effects of acupuncture manipulation and drugs, which is to inject drugs into relative acupoint or special point (Tang 2010). Laser acupuncture, guiding by the basic theory of Chinese medicine, is a treatment method of preventing disease, treating disease, and health care to stimulate acupoints effectively using low-intensity laser beam to irradiate acupoints directly, focus on acupoints or with beam expander (Fan 2010). Acupoint catgut treatment, guiding by the basic theory of Chinese medicine, is an external treatment method of treating diseases with catgut's stimulation effects on acupoints by using different types of catgut to embed acupoints selectively (Meng 2012). Electric acupuncture convulsive therapy, deriving from acupuncture, is a treatment method for mental disorders by using subconvulsive stimulating currents and with the electrodes in acupoints (Baihui and Renzhong) (Xue 1985).

#### 7.1 Acupuncture added to standard dose antipsychotics

#### 7.1.1 Acupuncture

The categories of acupuncture were traditional acupuncture (traditional - Bouhlel 2011; traditional - Liu 2010; traditional - Luo 2006; traditional - Ma 2008; traditional - Tang 2005); electroacupuncture (electro - Chen 2006; electro - Chen 2008; electro - Cheng 2009; electro - Ding 2005; electro - Wang 2005; electro - Yao 2006; electro - Zhang 1993; electro - Zhang 2001); acupoint injection (acupoint inj - Wang 2000; acupoint inj - Yang 2000); laser acupuncture (laser - Ma 1999); and acupoint catgut treatment (acupoint cat - Wang 1997). None of the above acupuncture interventions were the same.

#### 7.1.2 Antipsychotics

Five studies used risperidone and the dose of four of these studies (electro - Wang 2005; traditional - Liu 2010; traditional - Luo 2006; traditional - Ma 2008) ranged from 2 to 6 mg/d. electro - Cheng 2009 used the average dosage of  $5.15 \pm 0.46$  mg/d. One study, electro - Chen 2008, used aripiprazole (average dosage  $18.4 \pm 6.2$  mg/d); one, electro - Yao 2006, used clozapine (total dosage 200 to 300 mg/d); and one, laser - Ma 1999, chlorpromazine (average dosage 395

Acupuncture for schizophrenia (Review)

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



± 55 mg/d). The other three studies did not describe more details of antipsychotics (acupoint cat - Wang 1997; electro - Zhang 2001; traditional - Bouhlel 2011). The remaining six studies reported that people remained on previous antipsychotics treatment (acupoint inj - Wang 2000; acupoint inj - Yang 2000; electro - Chen 2006; electro - Ding 2005; electro - Zhang 1993; traditional - Tang 2005).

#### 7.2 Acupuncture added to low dose antipsychotics

# 7.2.1 Acupuncture

The categories of acupuncture were traditional acupuncture (traditional - Wang 2006; traditional - Xu 2004), electroacupuncture (electro - Cui 2000; electro - Xiong 2010; electro - Zhou 1997), acupoint injection (acupoint inj - Pan 2002), laser acupuncture (laser - Zhang 1991) and acupoint catgut treatment (acupoint cat - Sun 2005). None of them were identical.

#### 7.2.2 Low dose antipsychotics

Five studies (acupoint inj - Pan 2002; electro - Cui 2000; laser - Zhang 1991; traditional - Wang 2006; traditional - Xu 2004) used chlorpromazine less than 300 mg/d. One study, electro - Xiong 2010, reported the use of clozapine 100 to 150 mg/d, and another study, acupoint cat - Sun 2005, used risperidone 1 to 2 mg/d. Finally, electro - Zhou 1997 described the dosage of the antipsychotics was a reduction of ~60% of their previous daily levels.

#### 7.3 Acupuncture added to TCM drug

#### 7.3.1 Acupuncture

Two studies used traditional acupuncture (traditional - Zhao 2005a; traditional - Zhao 2005b) and one study, electro - Zhang 1987, used electroacupuncture.

#### 7.3.2 TCM drug

Two studies used Fuyuankang capsule (traditional - Zhao 2005a; traditional - Zhao 2005b) and one study, electro - Zhang 1987, traditional decoction of herbs - Dang Gui Cheng Qi Tang.

#### 7.4 Acupuncture

The categories of acupuncture were traditional acupuncture (traditional - Wang 2006; traditional - Zhao 2005a; traditional - Zhao 2005b), electroacupuncture (electro - Zhang 1987) and laser acupuncture (laser - Liu 1986; laser - Zhang 1991) with different acupoints.

#### 7.5 Antipsychotics

Eight studies used risperidone ranging from 2 to 10 mg/d (acupoint cat - Sun 2005; electro - Cheng 2009; electro - Wang 2005; traditional - Liu 2010; traditional - Luo 2006; traditional - Ma 2008; traditional - Zhao 2005a; traditional - Zhao 2005b). Six studies used chlorpromazine (acupoint inj - Pan 2002; electro - Cui 2000; electro - Zhang 1987; laser - Liu 1986; laser - Ma 1999; laser - Zhang 1991). Three studies did not describe more details of antipsychotics used (acupoint cat - Wang 1997; electro - Zhang 2001; traditional - Bouhlel 2011). Two studies reported using clozapine (electro -Xiong 2010; electro - Yao 2006), one study - electro - Chen 2008 - used aripiprazole and the average dosage was 20.1 ± 4.3 mg/ d. The remaining seven studies reported that people remained on previous antipsychotics treatment (acupoint inj - Wang 2000; acupoint inj - Yang 2000; electro - Chen 2006; electro - Ding 2005; electro - Zhang 1993; electro - Zhou 1997; traditional - Tang 2005). One study - traditional - Wang 2006 - reported using enough dosage antipsychotics, which was equal to chlorpromazine dosage ranged from 0.4 to 0.6 g/d) and traditional - Xu 2004 reported maximum daily dosage was equivalent chlorpromazine dosage 0.4 to 0.7 g/d.

# 7.6 TCM drug

Two studies used Fuyuankang capsule (traditional - Zhao 2005a; traditional - Zhao 2005b) and one - electro - Zhang 1987 - used traditional decoction of herbs - Dang Gui Cheng Qi Tang.

#### 7.7 Electric acupuncture convulsive therapy

Only one study, EACT - Xue 1987, reported using acupoints Renzhong and Baihui with average electricity consumption 1.27 Joule.

#### 7.8 Electric convulsive therapy

One study, EACT - Xue 1987, reported using classic electroconvulsive therapy with average electricity consumption 34.97 Joule.

#### 8. Outcomes

#### 8.1 General remarks

Most outcomes of global state, behaviour and adverse effects were dichotomous and trials used a variety of scales. Two studies also reported adding medication outcomes, see Table 1. However some of the scale-derived data were skewed data and difficult to understand.

#### 8.2 Outcomes scales from which it was possible to use data

#### 8.2.1 Global state scales

#### 8.2.1.1 Clinical Global Impression Scale - CGI (Guy 1970)

The CGI is a three-item scale commonly used in studies on schizophrenia that enables clinicians to quantify severity of illness and overall clinical improvement. The items are: severity of illness, global improvement and efficacy index. A seven-point scoring system is usually used with low scores indicating decreased severity and/or greater recovery. laser - Zhang 1991 and electro - Zhou 1997 reported CGI data.

#### 8.2.2 Mental state scales

8.2.2.1 Brief Psychiatric Rating Scale - BPRS (Overall 1962)

The BPRS is an 18-item scale measuring positive symptoms, general psychopathology and affective symptoms. The original scale has 16 items, but a revised 18-item scale is commonly used. Scores can range from zero to 126. Each item is rated on a seven-point scale, with high scores indicating more severe symptoms. Thirteen studies reported BPRS data (acupoint inj - Pan 2002; acupoint inj - Yang 2000; electro - Cui 2000; electro - Ding 2005; electro - Xiong 2010; electro - Zhang 1993; electro - Zhou 1997; laser - Ma 1999; laser - Zhang 1991; traditional - Liu 2010; traditional - Luo 2006; traditional - Wang 2006; traditional - Xu 2004). One study, acupoint inj - Wang 2000, only reported the data of the twelfth item of BPRS - hallucinations.

8.2.2.2 Positive and Negative Syndrome Scale - PANSS<sup>\*</sup> (Kay 1986) This is a 30-item scale, each of which can be defined on a seven point scoring system from absent to extreme. It has three subscales for measuring the severity of general psychopathology, positive symptoms (PANSS-P), and negative symptoms (PANSS-N). A low score indicates lesser severity. Eight studies reported data using this scale (acupoint cat - Sun 2005; electro - Chen 2008; electro

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



- Cheng 2009; electro - Ding 2005; electro - Wang 2005; electro - Yao 2006; traditional - Bouhlel 2011; traditional - Ma 2008).

# 8.2.2.3 Scale for the Assessment of Positive Symptoms - SAPS (Andreasen 1982)

This six-point scale gives a global rating of positive symptoms such as delusions, hallucinations and disordered thinking. Higher scores indicate more symptoms. Five studies reported SAPS data (acupoint cat - Sun 2005; electro - Zhang 1993; traditional - Bouhlel 2011; traditional - Wang 2006; traditional - Xu 2004).

8.2.2.4 Scale for the Assessment of Negative Symptoms - SANS (Andreasen 1982)

This scale allows a global rating of the following negative symptoms: alogia (impoverished thinking), affective blunting, avolition-apathy, anhedonia-asociality and attention impairment. Assessments are made on a six-point scale (zero = not at all to five = severe). Higher scores indicate more symptoms. Data for this scale were reported by five studies (electro - Zhang 1993; traditional - Bouhlel 2011; traditional - Luo 2006; traditional - Wang 2006; traditional - Xu 2004).

# 8.2.2.5 Hamilton Rating Scale for Depression - HAMD (Hamilton 1967)

This instrument is designed to be used only on patients already diagnosed as suffering from affective disorder of the depressive type. It is used for quantifying the results of an interview, and its value depends entirely on the skill of the interviewer in eliciting the necessary information. The scale contains 17 variables measured on either a five-point or a three-point rating scale, the latter being used where quantification of the variable is either difficult or impossible. Among the variables are: depressed mood, suicide, work and loss of interest, retardation, agitation, gastrointestinal symptoms, general somatic symptoms, hypochondriasis, loss of insight, and loss of weight. It is useful to have two raters independently scoring a patient at the same interview. The scores of the patient are obtained by summing the scores of the two physicians. A score of 11 is generally regarded as indicative of a diagnosis of mild depression, 14 to 17 mild to moderate depression and >17 moderate to severe depression. electro - Chen 2006 and electro - Zhang 2001 reported data of this scale.

#### 8.2.2.6 Zung Self-Rating Depression Scale - SDS (Zung 1965)

The Zung Self-Rating Depression Scale is a 20-item self-rated scale that is widely used as a screening tool, covering affective, psychological and somatic symptoms associated with depression. The questionnaire takes approximately 10 minutes to complete and items are framed in terms of positive and negative statements. It can be effectively used in a variety of settings, including primary care, psychiatric clinics, drug trials and various research situations. Each item is scored on a Likert scale ranging from one to four. Most people with depression score between 50 and 69, while a score of 70 and above indicates severe depression. electro - Zhang 2001 reported SDS data.

8.2.2.7 Psychotic symptom rating scale - PSYRHS<sup>\*</sup> (Haddock 1999) Psychotic Symptom Rating Scales is a 17-item, five-point scale to rate symptom scores (zero to four) with high scores indicating more severe symptoms. The scales consist of two subscales - the auditory hallucinations subscale (AH) and delusions subscale (DS). The auditory hallucinations subscale is an 11-item scale. The dimensions of auditory hallucinations are frequency, duration, location, loudness, beliefs re-origin of voices, amount of negative content of voices, degree of negative content, amount of distress, intensity of distress, disruption to life caused by voices, controllability of voices. The total AH score ranges from zero to 44. The delusions subscale is a six-item scale. The dimensions of delusions include amount of preoccupation with delusions, duration of preoccupation with delusions, conviction, amount of distress, intensity of distress and disruption to life caused by beliefs. The total DS score ranges from zero to 24. electro - Cheng 2009 reported data of auditory hallucinations subscale (PSYRHS-AH).

#### 8.2.2.8 Specific Auditory Hallucination Scale - SAHS<sup>\*</sup> (Lu 2006)

Specific Auditory Hallucination Scale derives from Scale for the Assessment of Positive Symptoms. The scale includes four items of SAPS. The four items are auditory hallucinations, voice commenting, voice conversing, global rating of severity of hallucinations. Higher scores indicate more symptoms. Only traditional - Liu 2010 reported SAHS data.

#### 8.2.3 Adverse effects scales

8.2.3.1 Treatment Emergent Symptom Scale/Form - TESS/F (Guy 1976)

This checklist assesses a variety of characteristics for each adverse event, including severity, relationship to the drug, temporal characteristics (timing after a dose, duration and pattern during the day), contributing factors, course and action taken to counteract the effect. Symptoms can be listed a priori or can be recorded as observed by the investigator. Fourteen studies used this scale (acupoint inj - Pan 2002; acupoint cat - Sun 2005; electro - Chen 2006; electro - Chen 2008; electro - Cui 2000; electro - Xiong 2010; electro - Yao 2006; electro - Zhang 1993; electro - Zhou 1997; traditional - Liu 2010; traditional - Ma 2008;electro - Wang 2005; traditional - Wang 2006; traditional - Xu 2004). electro - Zhang 1993 only reported part of data and only six studies reported scores of this scale (acupoint inj - Pan 2002; electro - Xiong 2010; electro -Yao 2006; electro - Zhou 1997; traditional - Wang 2006; traditional - Xu 2004), three of them with skewed data (electro - Xiong 2010; traditional - Wang 2006; traditional - Xu 2004).

8.2.3.2 Rating Scale for Extrapyramidal Side Effects - RESES\* (Simpson 1970)

Rating scale for Extrapyramidal Side Effects is a 10-item scale relating to extrapyramidal side effects. The score of each item rates symptom from zero to four. Zero means normal and high scores indicate severe side effects. The items are gait, arm dropping, shoulder shaking, elbow rigidity, wrist rigidity of fixation of position, pendulousness of legs, head dropping, glabella tap, tremor and salivation. Only one study, laser - Zhang 1991, used this scale and reported mean, events number of each group and T-value.

Outcomes with '\*' are extra outcomes added in this 2012 update.

#### 9. Missing outcomes

The included studies did not attempt to quantify outcomes of death, engagement with services, satisfaction with treatment, quality of life, economic outcomes.

#### **Excluded studies**

We excluded nine studies. Studies listed in the Characteristics of excluded studies had to be inspected in hard copy in order to make the final decision. One study (Sun 1994) was a case series.

Acupuncture for schizophrenia (Review)

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



Two studies, Ma 2004 and Zhuge 1993, were not randomised. Xiong 2009 was quasi-randomised. Participants of Zhong 1995 did not suffer from schizophrenia, schizophreniform psychosis or schizophrenia-like illnesses. Participants of Luo 2006a were people with schizophrenia or mood disorders manic episode and outcomes were not reported by diagnostic group. Two studies, Ma 2002 and Xue 1985, had to be excluded because of the interventions. One study (Wu 2004) was excluded because some important data reported in this study were identical to those of another included study (acupoint inj - Pan 2002) but the allocation numbers of the two groups, the acupoint choice method and the BPRS factor data were different. We could not acquire further details.

#### Awaiting assessment

One study (NCT01167348) awaits assessment as we have not acquired further details and were unable to confirm whether the trial has ended; we were unable to locate the full text of the study.

# Ongoing

We know of no ongoing studies.

# **Risk of bias in included studies**

We judged the risk of bias for the included studies and, overall, we judged that the risk of bias to be moderate (Figure 3; Figure 4).

# Figure 3. 'Risk of bias' graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.





Figure 4. 'Risk of bias' summary: review authors' judgements about each risk of bias item for each included study.



# Figure 4. (Continued)

laser - Ma 1999		?	?	?	+	?	•
laser - Zhang 1991	?	?	•	•	•	?	•
traditional - Bouhlel 2011	?	?	•	•		?	•
traditional - Liu 2010	•	?	•	?		?	?
traditional - Luo 2006	?	?	?	?	÷	?	•
traditional - Ma 2008	?	?	•	?	÷	?	•
traditional - Tang 2005	?	?	•	?	÷	?	•
traditional - Wang 2006	•	?	•	?	•	?	•
traditional - Xu 2004	•	?	•	?	÷	?	?
traditional - Zhao 2005a	?	?	•	?	•	•	•
traditional - Zhao 2005b	?	?	•	?	•	•	•

# Allocation

All 30 included studies were stated to be randomised, but only 10 provided descriptions of the randomisation method used (acupoint cat - Sun 2005; acupoint inj - Wang 2000; electro - Chen 2008; electro - Cheng 2009; electro - Cui 2000; electro - Yao 2006; electro - Zhang 2001; traditional - Liu 2010; traditional - Wang 2006; traditional - Xu 2004) and one (electro - Cheng 2009), also described allocation concealment. Two studies did not definitely report the number of participants in each group (electro - Zhang 1993; traditional - Bouhlel 2011). The number of participants in the remaining studies were evenly balanced in each group except acupoint inj - Pan 2002 (105/65), electro - Zhou 1997 (25/15), traditional - Zhao 2005a (90/90/90/30), and traditional - Zhao 2005b (90/90/90/30).

# Blinding

Sham acupuncture is difficult to use and it is unclear if this is a successful method though assessor blinding is easier. Two studies stated that both patients and assessors were blinded (electro - Cheng 2009; traditional - Bouhlel 2011). Five studies were assessor blinded (acupoint inj - Pan 2002; acupoint inj - Wang 2000; electro - Xiong 2010; electro - Zhou 1997; laser - Zhang 1991) and in one study, traditional - Liu 2010, it was with unclear whether it was single blind. The remaining 22 studies did not report whether assessor blinding had been used but participants of 19 trials were unblinded and three trials did not report if they used a method by which participants were blinded (acupoint cat - Wang 1997; laser - Ma 1999; traditional - Luo 2006).

# Incomplete outcome data

Five studies reported incomplete outcome data - totaling 71 participants (acupoint inj - Pan 2002; electro - Chen 2008; electro - Yao 2006; traditional - Bouhlel 2011; traditional - Liu 2010) and for the participants of one study, electro - Zhang 1993, it was unclear. Two studies used Intention-to-treat (ITT) method of analysis (electro - Cheng 2009; traditional - Xu 2004) and no participants of the other 23were lost to follow-up.

# Selective reporting

The protocols of 28 studies were not available, therefore we could not compare outcomes in the protocols with those in the published reports. Authors of another two studies recorded that BPRS was one of the outcomes but they only reported global state, rather than this measure (traditional - Zhao 2005a; traditional - Zhao 2005b).

# Other potential sources of bias

The references of two studies, acupoint inj - Pan 2002 (supported by government) and electro - Zhang 1993, reported different data and one of the authors of acupoint inj - Wang 2000 worked for the drug industry, hence these three studies have potential sources of bias. Three studies were supported by government or an institute (acupoint cat - Sun 2005; electro - Cheng 2009; traditional - Liu 2010). Two studies (electro - Ding 2005; traditional - Xu 2004), only included men. We were unclear whether these five studies had 'other potential sources' of bias. Overall, 29 of 30 included studies in this update were from the People's Republic of China. It is unclear if this represents a racial or cultural bias when applied to other regions.

# **Effects of interventions**

See: Summary of findings for the main comparison ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia; Summary of findings 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia; Summary of findings 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS for schizophrenia; Summary of findings 4 ACUPUNCTURE added to TCM DRUG versus TCM DRUG for schizophrenia; Summary of findings 5 ACUPUNCTURE versus TCM DRUG for schizophrenia; Summary of findings 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY for schizophrenia

It is impossible to judge if different categories of acupuncture really have different mechanisms of action. We undertook subgroup

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd. analyses to investigate effects for the same outcome of various categories of acupuncture and, when heterogeneity was low, we synthesised data. We also added a table of adverse effects of acupuncture according to different categories (Table 2).

For the previous version of description of studies, please see Appendix 5.

We calculated risk ratios (RR) for dichotomous data and estimated mean differences (MD) for continuous data, with their respective 95% confidence intervals (CIs) throughout.

#### 1. COMPARISON 1: ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Seventeen studies (acupoint cat - Wang 1997; acupoint inj - Wang 2000; acupoint inj - Yang 2000; electro - Chen 2006; electro - Chen 2008; electro - Cheng 2009; electro - Ding 2005; electro - Wang 2005; electro - Yao 2006; electro - Zhang 1993; electro - Zhang 2001; laser - Ma 1999; traditional - Liu 2010; traditional - Luo 2006; traditional - Ma 2008; traditional - Tang 2005; traditional - Bouhlel 2011) compared acupuncture added to standard dose antipsychotics with standard dose antipsychotics.

#### 1.1 Global state

# 1.1.1 Not improved

The binary data of three studies with traditional acupuncture used three different criteria of 'not improved' at medium-term but the criteria were similar. We found medium-term global state data favoured acupuncture added to standard dose antipsychotics (n = 244, 3 RCTs, RR 0.40 CI 0.28 to 0.57). We, however, found a different result from one study with laser acupuncture where the criteria and duration of study were unclear (n = 120, 1 RCT, RR 0.20 CI 0.01 to 4.08, Analysis 1.1).

# 1.1.2 Adding medication

Only electro - Wang 2005 reported using additional medication, which was not a pre-stated outcome of interest for this review, but we have listed these data in Table 1.

# 1.2 Mental state

#### 1.2.1 Brief Psychiatric Rating Scale (BPRS) endpoint score

We found short-term BPRS scores of four studies - two with traditional acupuncture, one with electroacupuncture and one with laser acupuncture - significantly favoured acupuncture added to standard dose antipsychotics compared with standard dose antipsychotics alone (n = 327, 4 RCTs, MD -4.32 CI -5.28 to -3.36). Three studies reported medium-term BPRS scores with considerable heterogeneity  $(1^2 > 75\%, Analysis 1.2)$ . With the medium-term subgroup analysis, according to different categories of acupuncture, we found a substantial heterogeneity between traditional acupuncture and the acupoint injection subgroup (12 > 50%) and also between traditional - Liu 2010 and traditional - Luo 2006 in the same subgroup ( $l^2 > 75\%$ ). With traditional acupuncture we found that those given traditional acupuncture added to standard dose antipsychotics had a significantly better rating in their mental state compared with the standard dose antipsychotics group (traditional - Liu 2010: n = 96, 1 RCT, MD -4.87 CI -16.24 to -3.50; traditional - Luo 2006: n = 60, 1 RCT, MD -11.35 CI -14.48 to -8.22). However, there was no difference between acupoint injection added to standard dose antipsychotics and standard dose antipsychotics alone (n = 64, 1 RCT, MD -0.70 CI -5.02 to 3.62, Analysis 1.3)

#### 1.2.2 Positive and Negative Syndrome Scale (PANSS) endpoint score

One study with traditional acupuncture and three studies with electroacupuncture reported short-term PANSS scores without skewed data. Data indicated considerable heterogeneity (I<sup>2</sup> > 75%) but, by the medium-term, there was significant difference between acupuncture added to standard dose antipsychotics and standard dose antipsychotics alone group (n = 135, 2 RCTs, MD -3.79 CI -6.43 to -1.15, Analysis 1.4). With subgroup analysis, we found there was a considerable heterogeneity between these two categories of acupuncture (I<sup>2</sup> > 75%). With traditional acupuncture there was only one study and this indicated no difference between traditional acupuncture added to standard dose antipsychotics and standard dose antipsychotics alone (n = 60, 1 RCT, MD 1.30 CI -1.56 to 4.16).

The PANSS endpoint score from the three studies with electroacupuncture contained substantial heterogeneity ( $I^2 > 50\%$ , Analysis 1.5). Removing the study with results that were causing this heterogeneity, as judged by visual inspection (electro Ding 2005), eliminated this heterogeneity. We found short-term data favoured electroacupuncture added to the standard dose antipsychotics group (n = 135, 2 RCTs, MD -3.79 Cl -6.43 to -1.15, Analysis 1.6). Three studies also used binary data to assess PANSS at short-term with the assertion that less than 25% reduction as 'not improved' (electro - Chen 2008; electro - Wang 2005; traditional -Ma 2008). There was a difference between acupuncture added to standard dose antipsychotics and standard dose antipsychotics (n = 197, 3 RCTs, RR 0.65 CI 0.45 to 0.94). The electro - Yao 2006 trial used binary data to assess PANSS at short-term and medium-term - with assumption of less than 30% reduction as 'not improved'. There was no difference between electroacupuncture added to standard dose antipsychotics and standard dose antipsychotics alone in the short-term (n = 90, 1 RCT, RR 0.91 CI 0.43 to 1.92) or by medium-term (n = 90, 1 RCT, RR 0.69 CI 0.36 to 1.31, Analysis 1.8). Three studies reported skewed PANSS data at short-term and one study provided skewed PANSS data at medium-term (Analysis 1.7).

# 1.2.3 Scale for the Assessment of Positive Symptoms (SAPS) endpoint score

Two studies, one with traditional acupuncture and one with electroacupuncture, reported SAPS scores (short-term) but these were skewed continuous data (Analysis 1.9).

# 1.2.4 Scale for the Assessment of Negative Symptoms (SANS) endpoint score

The traditional - Luo 2006 trial used the SANS to assess mental state. Those given traditional acupuncture added to standard dose antipsychotics had a significantly better rating in their negative symptoms compared with the standard dose antipsychotics group (short-term, n = 60, 1 RCT, MD -7.66 CI -13.05 to -2.27; medium-term, n = 60, 1 RCT, MD -12.35 CI -17.54 to -7.16, Analysis 1.10). Another two studies reported short-term skewed SANS endpoint score data (Analysis 1.11).

#### 1.2.5 Hamilton Rating Scale for Depression (HAMD) endpoint score

Only two studies with electroacupuncture used this scale for short-term assessment. There was a considerable heterogeneity between these two studies ( $I^2 = 75\%$ ). We found the scores of electroacupuncture added to the standard dose antipsychotics

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



group were lower than the standard dose antipsychotics group (electro - Chen 2006: n = 67, 1 RCT, MD - 6.90 Cl - 9.31 to -4.49; electro - Zhang 2001: <math>n = 42, 1 RCT, MD - 10.41 Cl - 12.81 to -8.01, Analysis 1.12). A similar result was found if converting continuous data to binary data but heterogeneity disappeared (n = 109, 2 RCTs, RR 0.17 Cl 0.08 to 0.34,  $l^2 = 0\%$ , Analysis 1.13).

#### 1.2.6 Self-Rating Depression Scale (SDS) endpoint score

Continuous data of the only study to report relevant data indicated a significant difference between electroacupuncture added to standard dose antipsychotics compared with standard dose antipsychotics alone (short-term, n = 42, 1 RCT, MD -24.25 Cl -28.01 to -20.49, Analysis 1.16).

#### 1.2.7 Psychotic Symptom Rating Scales Auditory Hallucination Subscale (PSYRAS-AH) endpoint score

We found only one study, electro - Cheng 2009, which used the PSYRAS-AH subscale to assess specific symptoms. Participants given electroacupuncture added to standard dose antipsychotics had a lower score than those only given standard dose antipsychotics alone (short-term, n = 60, 1 RCT, MD -2.17 CI -4.16 to -0.18, Analysis 1.15). Converting continuous data to binary found a similar result (n = 60, 1 RCT, RR 0.27 CI 0.14 to 0.52, Analysis 1.14).

#### 1.2.8 Auditory hallucinations not improved

acupoint cat - Wang 1997, acupoint inj - Yang 2000 and laser - Ma 1999 reported whether people heard auditory hallucinations at different time periods using different criteria for 'not improved'. We found a significant difference between acupuncture added to standard dose antipsychotics group compared with the standard dose antipsychotics group when acupoint catgut treatment was used (short-term, n = 216, 1 RCT, RR 0.24 CI 0.13 to 0.44) or acupoint injection (medium-term, n = 64, 1 RCT, RR 0.32 CI 0.17 to 0.61), but not when comparing laser acupuncture added to standard dose antipsychotics with standard dose antipsychotics alone (n = 120, 1 RCT, RR 0.25 CI 0.03 to 2.17), though the duration of this study was unclear (Analysis 1.17).

#### 1.2.9 Specific Auditory Hallucination Scale (SAHS) endpoint score

traditional - Liu 2010 reported skewed continuous data from SAHS for the short- and medium-term (Analysis 1.18).

#### 1.2.10 Hallucinations

acupoint inj - Wang 2000 reported data from the twelfth item of the BPRS. We found short-term data were significantly different between acupoint injection added to standard dose antipsychotics and standard dose antipsychotics alone (n = 90, 1 RCT, MD -0.73 CI -1.28 to -0.18, Analysis 1.19).

#### 1.2.11 Time for auditory hallucinations to disappear

laser - Ma 1999 reported this outcome. These data are listed using additional Table 1.

#### 1.3 Behaviour

#### 1.3.1 Leaving the study early

We found 10 studies with five different categories of acupuncture (all short-term data). Only seven of 870 participants left the studies early. There was no difference between acupuncture added to standard dose antipsychotics group and standard dose antipsychotics alone (n = 870, 10 RCTs, RR 1.33 CI 0.33 to

Acupuncture for schizophrenia (Review)

Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

5.45). Five of 370 participants with either acupoint injection, electroacupuncture or traditional acupuncture left the studies early by the medium-term (n = 370, 5 RCTs, RR 3.58 CI 0.60 to 21.27, Analysis 1.20).

#### **1.4 Service outcomes**

#### 1.4.1 Time in hospital

Only one study with laser acupuncture reported the outcome of time in hospital. Although this was not a pre-stated outcome of interest for this review, it was the only service outcome and we analysed data and found the days in hospital of participants given laser acupuncture added to standard dose antipsychotics was less than the number of days of participants given standard dose antipsychotics alone (n = 120, 1 RCT, MD -16.00 CI -19.54 to -12.46, Analysis 1.21).

# 1.5 Adverse effects

# 1.5.1 Treatment Emergent Symptom Scale/Form (TESS) endpoint score

electro - Yao 2006 reported TESS endpoint score at short-term with result significantly favouring the electroacupuncture added to standard dose antipsychotics group (n = 90, 1 RCT, MD -2.80 CI -3.09 to -2.51, Analysis 1.22).

#### 1.5.2 Extrapyramidal symptoms

Acupuncture added to standard dose antipsychotics did not reduce the incidence of extrapyramidal symptoms in the shortterm (n = 202, 3 RCTs, RR 0.66 CI 0.40 to 1.09). There was substantial heterogeneity between traditional acupuncture and electroacupuncture group (traditional acupuncture: n = 60, 1 RCT, RR 0.17 CI 0.02 to 1.30; electroacupuncture: n = 142, 2 RCTs, RR 0.79 CI 0.47 to 1.34, subgroup I<sup>2</sup>>50%). We found a similar result at medium-term (n = 156, 2 RCTs, RR 0.62 CI 0.32 to 1.23). Further, electro - Wang 2005 reported there was no difference of the rate of tremor (n = 75, 1 RCT, RR 0.73 CI 0.24 to 2.18) and akathisia (n = 75, 1 RCT, RR 0.58 CI 0.23 to 1.48) if electroacupuncture was added to standard dose antipsychotics group compared with standard dose antipsychotics alone in the short-term. electro - Chen 2008 reported no difference in rate of myotonia (n = 60, 1 RCT, RR 1.00 CI 0.07 to 15.26), tremor (n = 60, 1 RCT, RR 0.67 CI 0.12 to 3.71) and akathisia (n = 60, 1 RCT, RR 0.60 CI 0.16 to 2.29) when electroacupuncture was added to standard dose antipsychotics group compared with standard dose antipsychotics alone by the medium-term (Analysis 1.23; Analysis 1.24).

#### 1.5.3 Central Nervous System

traditional - Liu 2010 provided incidences of anxiety in the mediumterm and we found traditional acupuncture added to standard dose antipsychotics did not reduce the incidences compared with standard dose antipsychotics alone (n = 96, 1 RCT, RR 0.42 Cl 0.09 to 2.05). Data from electro - Chen 2006, electro - Wang 2005 or traditional - Ma 2008 indicated acupuncture added to standard dose antipsychotics reduced the frequency of insomnia compared with standard dose antipsychotics by the short-term (n = 202, 3 RCTs, RR 0.30 Cl 0.11 to 0.83), however, two studies reported no difference by the medium-term (n = 156, 2 RCTs, RR 0.34 Cl 0.12 to 1.02, Analysis 1.25). electro - Chen 2008 reported no different for headache between electroacupuncture added to standard dose antipsychotics group compared with standard dose antipsychotics alone by the medium-term (n = 60, 1 RCT, RR 3.00 CI 0.33 to 27.23, Analysis 1.25).

### 1.5.4 Anticholinergic symptoms

We found no difference in dry mouth in the short-term (n = 202, 3 RCTs, RR 1.18 CI 0.47 to 3.00), blurred vision (n = 202, 3 RCTs, short-term RR 0.88 CI 0.24 to 3.24; n = 156, 2 RCTs, medium-term RR 1.00 CI 0.15 to 6.64), sweating (n = 156, 2 RCTs, medium-term RR 3.00 CI 0.13 to 70.83), constipation (n = 202, 3 RCTs, short-term RR 0.88 CI 0.19 to 4.06) and nausea and vomiting (n = 202, 3 RCTs, short-term RR 0.41 CI 0.12 to 1.32) when comparing acupuncture added to standard dose antipsychotics and standard dose antipsychotics alone (Analysis 1.26).

# 1.5.5 Gastrointestinal system

People given traditional acupuncture added to standard dose antipsychotics had similar 'unspecified gastrointestinal symptoms' by the medium-term (n = 96, 1 RCT, RR 0.52 CI 0.10 to 2.71), constipation (n = 202, 3 RCTs, RR 0.88 CI 0.19 to 4.06) and nausea and vomiting (n = 202, 3 RCTs, RR 0.41 CI 0.12 to 1.32) in the short-term as those only given standard dose antipsychotics alone (Analysis 1.27).

# 1.5.6 Cardiovascular symptoms

Acupuncture added to standard dose antipsychotics group did not reduce dizziness (n = 202, 3 RCTs, short-term RR 0.94 CI 0.32 to 2.75), dizziness or headache (n = 96, 1 RCT, medium-term RR 0.35 CI 0.04 to 3.22), or tachycardia (n = 217, 3 RCTs, short-term RR 0.87 CI 0.42 to 1.79; n = 156, 2 RCTs, medium-term RR 0.34 CI 0.04 to 3.20, Analysis 1.28).

#### 1.5.7 Metabolic system

Comparing acupuncture added to standard dose antipsychotics group with standard dose antipsychotics group indicated substantial heterogeneity (I<sup>2</sup> > 50%) but when undertaking subgroup analysis we found acupuncture added to standard dose antipsychotics did not change weight gain compared with standard dose antipsychotics alone in the short-term - traditional acupuncture (n = 60, 1 RCT, RR 0.14 CI 0.01 to 2.65) and electroacupuncture (n = 142, 2 RCTs, RR 6.15 CI 0.33 to 115.01). The same held for the findings for traditional acupuncture in the medium-term (n = 96, 1 RCT, RR 0.21 CI 0.01 to 4.23, Analysis 1.29; Analysis 1.30).

#### 1.5.8 Endocrine system

There was no difference in the rate of irregular menstruation (short-term) between acupuncture added to standard dose antipsychotics group and standard dose antipsychotics alone (n = 127, 2 RCTs, RR 0.33 Cl 0.01 to 7.87, Analysis 1.31).

#### 1.5.9 Laboratory tests

Short-term (n = 292, 4 RCTs, RR 1.01 CI 0.26 to 3.90) and mediumterm (n = 96, 1 RCT, RR 0.70 CI 0.12 to 3.98) data indicate acupuncture added to standard dose antipsychotics did not affect the rate of abnormal liver function tests. This also applied to ECG abnormalities (n = 292, 4 RCTs, RR 0.50 CI 0.05 to 5.32). Overall, participants allocated to acupuncture had no different rates of blood abnormalities in routine tests than those given only standard dose antipsychotics (n = 292, 4 RCTs, RR 1.33 CI 0.32 to 5.62, Analysis 1.32).

# 2. COMPARISON 2: ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Eight studies (acupoint cat - Sun 2005; acupoint inj - Pan 2002; electro - Cui 2000; electro - Xiong 2010; electro - Zhou 1997; laser - Zhang 1991; traditional - Wang 2006; traditional - Xu 2004) compared acupuncture added to low dose antipsychotics with standard dose antipsychotics.

# 2.1 Global state

# 2.1.1 Relapse

Only one study (acupoint inj - Pan 2002) with acupoint injection reported binary data for 'relapse'. We found acupoint injection added to low dose antipsychotics group reduced the rate of relapse compared with standard dose antipsychotics alone at long-term follow-up (n = 170, 1 RCT, RR 0.57 Cl 0.37 to 0.89, Analysis 2.1).

#### 2.1.2 Not improved

Four studies (acupoint cat - Sun 2005; electro - Zhou 1997; laser - Zhang 1991; traditional - Wang 2006), with traditional acupuncture, electroacupuncture, laser acupuncture or acupoint catgut treatment, used three different criteria of 'not improved' (all short-term). We found global data did not significantly favour acupuncture added to low dose antipsychotics (n = 272, 4 RCTs, RR 0.83 Cl 0.40 to 1.72, Analysis 2.2).

#### 2.1.3 Clinical Global Impression Scale (CGI) endpoint score

The CGI consists of three item - severity of illness (SI), global improvement (GI) and efficacy index (EI). electro - Zhou 1997 reported continuous data (short-term) but data of CGI-GI and CGI-EI were skewed (Analysis 2.4; Analysis 2.5). We found no difference of CGI-SI at short-term between participants given electroacupuncture added to low dose antipsychotics and those given standard dose antipsychotics (n = 40, 1 RCT, MD -0.40 CI -1.08 to 0.28, Analysis 2.3).

#### 2.1.4 Adding medication

One study (electro - Cui 2000) reported using adding medication, which was not the outcome we measured, but we listed it using additional table (Table 1).

# 2.2 Mental state

#### 2.2.1 BPRS endpoint/change score

We found continuous data from BPRS for the short-term were not significantly different between acupuncture added to low dose antipsychotics group and standard dose antipsychotics alone group but with considerable heterogeneity between each study (n = 332, 5 RCTs, MD -5.55 CI -14.40 to 3.29; I<sup>2</sup> > 75%). If we removed the study that was causing this heterogeneity, as judged by visual inspection (traditional - Wang 2006) and undertook a subgroup analysis, we found a similar conclusion (n = 300, 4 RCTs, MD -1.36 CI -3.13 to 0.41,  $I^2 = 49\%$ , subgroup  $I^2 = 49.3\%$ ). However, it was different for the long-term follow-up (n = 137, 1 RCT, MD -4.87 CI -8.21 to -1.53, Analysis 2.6; Analysis 2.7). Three studies also used binary data to assess BPRS (short-term) with no difference between acupuncture added to low dose antipsychotics and standard dose antipsychotics alone even with different reduction of BPRS as criteria (reduced rate < 30%: n = 60, 1 RCT, RR 0.75 CI 0.18 to 3.07; reduced rate < 25%: n = 170, 1 RCT, RR 0.62 CI 0.31 to 1.25; reduced rate ≤ 20%: n = 40, 1 RCT, RR 0.90 CI 0.30 to 2.68, Analysis 2.8). Only

Acupuncture for schizophrenia (Review)

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

one study reported BPRS change scores (short-term) and we found there was no difference between electroacupuncture added to low dose antipsychotics group and standard dose antipsychotics alone group (n = 60, 1 RCT, MD 0.81 CI -3.17 to 4.79, Analysis 2.9).

#### 2.2.2 PANSS endpoint score

The PANSS was used to assess symptoms by acupoint cat - Sun 2005 and electro - Xiong 2010. Data are skewed (Analysis 2.10). electro - Xiong 2010 also used binary data to assess PANSS and found no difference between electroacupuncture added to low dose antipsychotics and standard dose alone antipsychotics (n = 80, 1 RCT, RR 1.22 Cl 0.57 to 2.62, Analysis 2.11).

# 2.2.3 SAPS endpoint score

Three studies (acupoint cat - Sun 2005; traditional - Wang 2006; traditional - Xu 2004) with traditional acupuncture or acupoint catgut treatment added to low dose antipsychotics reported this result in the short-term. One study (traditional - Xu 2004). reports skewed data (Analysis 2.13).The other two, using different categories, had considerable heterogeneity ( $l^2 > 75\%$ ). Data from traditional - Wang 2006 indicated significant difference between traditional acupuncture added to low dose antipsychotics group and standard dose antipsychotics group (n = 32, 1 RCT, MD -13.34 Cl -16.94 to -9.74) but there was no difference between acupoint catgut treatment added to low dose antipsychotics group and standard dose antipsychotics alone group (n = 180, 1 RCT, MD 1.21 Cl 0.96 to 1.46, Analysis 2.12).

# 2.2.4 SANS endpoint score

traditional - Xu 2004 reported no difference for SANS endpoint score (short-term) between traditional acupuncture added to low dose antipsychotics group and standard dose antipsychotics alone group (n = 80, 1 RCT, MD 0.61 Cl -3.30 to 4.52, Analysis 2.14). traditional - Wang 2006 also reported data from SANS endpoint score but the data were skewed (Analysis 2.15).

# 2.3 Behaviour

# 2.3.1 Leaving the study early

We found participants given acupuncture added to low dose antipsychotics were not less likely to drop out than those give standard dose antipsychotics at short-term (n = 662, 8 RCT, RR 0.81 CI 0.29 to 2.29, Analysis 2.16).

# 2.4 Adverse effects

# 2.4.1 TESS endpoint score

Five studies (acupoint inj - Pan 2002; electro - Xiong 2010; electro - Zhou 1997; traditional - Wang 2006;traditional - Xu 2004) reported short-term TESS endpoint score but three sets of data were skewed (Analysis 2.18). The remaining two indicated significant differences between acupuncture added to low dose antipsychotics group and standard dose antipsychotics group (n = 200, 2 RCTs, MD -0.56 CI -0.86 to -0.26, Analysis 2.17).

# 2.4.2 Rating Scale for Extrapyramidal Side Effects (RESES) endpoint score

One study (laser - Zhang 1991) reported RESES endpoint score but only reported mean and T-value. We calculated SD according to the *Cochrane Handbook for Systematic Reviews of Interventions* ( Higgins 2011) and data indicated that scores of acupuncture added

**Cochrane** Database of Systematic Reviews

to low dose antipsychotics group were lower than those in the standard dose antipsychotics group (n = 20, 1 RCT, MD -0.60 CI -0.73 to -0.47, Analysis 2.19).

#### 2.4.3 Extrapyramidal symptoms

Studies with three categories of acupuncture had considerable heterogeneity between subgroups in the short-term (I<sup>2</sup> > 75%). The incidences of extrapyramidal symptoms were less for those in the acupuncture group whether given acupoint catgut (n = 180, 1 RCT, RR 0.02 CI 0.00 to 0.35) or electroacupuncture (n = 60, 1 RCT, RR 0.36 CI 0.15 to 0.87) - both added to low dose antipsychotics. Laser acupuncture, however, seemed not to make any difference in one small trial (n = 20, 1 RCT, RR 0.88 CI 0.53 to 1.46). There were no differences for short-term incidences of myotonia (n = 180, 1 RCT, RR 0.21 CI 0.01 to 4.29) and tremor (n = 180, 1 RCT, RR 0.09 CI 0.01 to 1.69) but data indicated acupoint catgut treatment added to low dose antipsychotics group (short-term, n = 180, 1 RCT, RR 0.03 CI 0.00 to 0.49, Analysis 2.20)

# 2.4.4 Central Nervous System

acupoint cat - Sun 2005 reported "activity-increasing adverse effects". We found acupoint catgut treatment added to low dose antipsychotics group could not reduce the event rate compared with standard antipsychotics group in the short-term (n = 180, 1 RCT, RR 0.15 Cl 0.01 to 2.85). acupoint cat - Sun 2005 also reported short-term insomnia. There was no difference for insomnia between acupoint catgut treatment added to low dose antipsychotics group and standard dose antipsychotics alone (n = 180, 1 RCT, RR 0.07 Cl 0.00 to 1.20). We did not find a clear difference for sleeplessness (n = 240, 2 RCTs, RR 0.33 Cl 0.04 to 3.03, Analysis 2.21)

#### 2.4.5 Anticholinergic symptoms

In the short-term, data for dry mouth favoured acupuncture (n = 240, 2 RCTs, RR 0.22 CI 0.09 to 0.58). Nasal congestion (short-term) was no different between groups (n = 180, 1 RCT, RR 0.09 CI 0.01 to 1.69). Two studies indicated that acupuncture added to low dose antipsychotics could reduce blurred vision (short-term, n = 240, 2 RCTs, RR 0.22 CI 0.06 to 0.83). There was no difference in constipation (n = 180, 1 RCT, short-term RR 0.07 CI 0.00 to 1.20, Analysis 2.22)

# 2.4.6 Cardiovascular symptoms

Short-term data indicate that there was no difference for dizziness between the two groups - acupoint catgut treatment added to low dose antipsychotics group and standard dose antipsychotics alone group (n = 180, 1 RCT, RR 0.70 CI 0.12 to 4.07). We found participants given acupuncture added to low dose antipsychotics had less tachycardia than people given standard dose antipsychotics alone (n = 240, 2 RCTs, RR 0.25 CI 0.11 to 0.53, Analysis 2.23).

# 2.4.7 Skin infection

No study found skin infection (Analysis 2.24).

# 3. COMPARISON 3: ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS

Six studies (electro - Zhang 1987; laser - Liu 1986; laser - Zhang 1991; traditional - Wang 2006; traditional - Zhao 2005a; traditional - Zhao 2005b) compared acupuncture with standard dose antipsychotics.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



# 3.1Global state

#### 3.1.1 Not improved

Five studies (electro - Zhang 1987; laser - Zhang 1991; traditional - Wang 2006; traditional - Zhao 2005a; traditional - Zhao 2005b) with a duration of less than three months are included in this comparison. Three (traditional - Wang 2006; traditional - Zhao 2005b) used traditional acupuncture, one electroacupuncture and the other laser acupuncture. Although studies all reported 'clinically important change' in global state, the criteria of 'not improved' were different except for traditional - Zhao 2005a and traditional - Zhao 2005b. We found no difference between acupuncture and standard dose antipsychotics which used similar criteria (n = 141, 3 RCTs, RR 0.88 CI 0.53 to 1.48). However, the other two trials, traditional - Zhao 2005a and traditional - Zhao 2005b, with identical criteria, indicated a difference between two groups (n = 240, 2 RCTs, RR 0.33 CI 0.19 to 0.59, Analysis 3.1)

# 3.2 Mental state

All included studies were with short-term.

#### 3.2.1 BPRS endpoint score

Only two studies reported BPRS. Continuous data favoured traditional acupuncture (n = 32, 1 RCT, MD -11.56 CI -16.36 to -6.76) but other continuous data indicated no difference between laser acupuncture and standard dose antipsychotics (n = 21, 1 RCT, MD -1.07 CI -11.19 to 9.05). There is substantial heterogeneity between two studies (I<sup>2</sup> > 50%, Analysis 3.2).

#### 3.2.2 SAPS endpoint score

Only traditional - Wang 2006 reported SAPS. We found scores of participants given traditional acupuncture were significantly less than those given antipsychotics (n = 32, 1 RCT, MD -5.24 Cl -9.06 to -1.42, Analysis 3.3).

#### 3.2.3 SANS endpoint score

traditional - Wang 2006 indicated no difference between traditional acupuncture and antipsychotics (n = 32, 1 RCT, MD -7.92 CI -17.01 to 1.17, Analysis 3.4).

# 3.3 Behaviour

We included six studies and there were no participants who left the studies early in each group in the short-term (Analysis 3.5).

# 3.4 Adverse effects

#### 3.4.1 TESS endpoint score

traditional - Wang 2006 reported TESS scores at short-term, but data were skewed and problematic to present (Analysis 3.6).

# 3.4.2 Extrapytamidal symptoms

Extrapyramidal symptoms (short-term) were lower in the laser acupuncture group with no participants experiencing this problem but eight out of 10 people reported this in the control group (n = 21, 1 RCT, RR 0.05 Cl 0.00 to 0.83, Analysis 3.7).

# 3.4.3 Numbness over the ear, upper extremity and chest on the treated side

We found no difference in numbness over the ear, upper extremity and chest on the treated side between participants given laser acupuncture and those given standard dose antipsychotics alone (n = 40, 1 RCT, RR 3.00 CI 0.13 to 69.52, Analysis 3.8).

# 4. COMPARISON 4: ACUPUNCTURE added to TCM DRUG versus TCM DRUG

Three studies (laser - Liu 1986; traditional - Zhao 2005a; traditional - Zhao 2005b) compared acupuncture added to TCM drug with TCM drug.

# 4.1 Global state

# 4.1.1 Not improved

We found a significant difference for short-term 'clinically important change' in global state between traditional acupuncture added to the TCM drug group and TCM drug group alone by studies with the same acupuncture criteria (n = 360, 2 RCTs, RR 0.11 CI 0.02 to 0.59), but no difference between electroacupuncture added to TCM drug group and TCM drug group alone by electro - Zhang 1987 (n = 94, 1 RCT, RR 0.77 CI 0.57 to 1.06, Analysis 4.1).

# 4.2 Behaviour

#### 4.2.1 Leaving the study early

We found no participants left the study early in either group (Analysis 4.2).

# 5. COMPARISON 5: ACUPUNCTURE versus TCM DRUG

Three studies (electro - Zhang 1987; traditional - Zhao 2005a; traditional - Zhao 2005b) compared acupuncture with TCM drug.

# 5.1 Global state

#### 5.1.1 Not improved

Three studies (electro - Zhang 1987; traditional - Zhao 2005a; traditional - Zhao 2005b) used two different criteria of 'not improved'. Two studies (traditional - Zhao 2005a; traditional - Zhao 2005b) with the same criteria indicated that there was no significant difference between traditional acupuncture group and TCM drug group in the short-term (n = 360, 2 RCTs, RR 1.46 CI 0.74 to 2.87). However, we found that participants given electroacupuncture were significantly less likely to experience a worsening in global state (n = 88, 1 RCT, RR 0.52 CI 0.34 to 0.80, Analysis 5.1).

#### 5.2 Behaviour

#### 5.2.1 Leaving the study early

electro - Zhang 1987, traditional - Zhao 2005a and traditional - Zhao 2005b reported data and we found no participant left the studies early in either acupuncture group or TCM drug group at short-term (Analysis 5.2).

# 6. COMPARISION 6: ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTRIC CONVULSIVE THERAPY

Only one study (EACT - Xue 1987) compared electric acupuncture convulsive therapy with electroconvulsive therapy at short-term.



# 6.1 Behaviour

#### 6.1.1 Leaving the study early

No participants left the study early (Analysis 6.1).

# 6.2 Adverse effects

#### 6.2.1 Back pain

EACT - Xue 1987 indicated there was no difference in the rates of back pain between electric acupuncture convulsive therapy group and standard electroconvulsive therapy group (n = 68, 1 RCT, RR 0.67 Cl 0.12 to 3.74, Analysis 6.2)

# 6.2.2 Spinal fracture

Unlike back pain, the same study indicated that electric acupuncture convulsive therapy causes lower incidences of spinal fracture than electroconvulsive therapy (n = 68, 1 RCT, RR 0.33 CI 0.14 to 0.81, Analysis 6.3).

# DISCUSSION

# Summary of main results

The main results are presented in 'Summary of findings' tables. Thirty studies were included and all findings are, at the very least, at moderate risk of bias. Ten trials describe the randomisation method but only one allocation concealment (by envelopes). Eight studies used a blinding method. Most studies were short-term and only two followed up participants to beyond six months. All but one study was undertaken in China.

# 1. Diagnosis and assessment criteria

Operational criteria (CCMD, ICD-10, DSM) were used in most studies although three trials did not report criteria. The recent version of the CCMD is CCMD-3. Whilst this refers to ICD-10 and the DSM-IV, it contains locally salient features that are absent in the international systems (Chen 2002). Many Chinese psychiatrists consider that this system has advantages with its inclusion of culture-distinctive categories and the exclusion of irrelevant Western diagnostic categories (Lee 2001). The differences between local and global psychiatric classifications causes some questions about diagnosis and assessment (Lee 2009). We are reasonably confident, however, that those entering these trials would be widely recognised as suffering from schizophrenia and even if 10% of diagnoses were inaccurate, this may reflect the real world more closely than if all diagnoses were entirely accurate.

We found that different criteria for outcomes of dichotomous data from global state and mental state made it difficult to merge data and analyse. We do think there is a strong argument for rationalising outcome measures (COMET).

# 2. Different treatments and controls

# 2.1 Acupuncture methods

With the development of acupuncture and modern medical technology, there are many categories of acupuncture. In this update review we found six acupuncture categories - traditional acupuncture (nine studies), electroacupuncture (12 studies), acupoint injection (three studies), laser acupuncture (three studies), acupoint catgut treatment (two studies) and another special category electric acupuncture convulsive therapy (one study). The mechanisms of action of any of these techniques

still could not be explained. Even with the same category of acupuncture there are various choices for acupoints, frequencies and durations (Table 3). It remains impossible to pre-judge clinical or physiological heterogeneity - or homogeneity - of different categories of acupuncture.

# 2.2 Placebo methods

In this update we found traditional - Bouhlel 2011 used sham traditional acupuncture, electro - Cheng 2009 used sham electroacupuncture, laser - Liu 1986 used sham laser irradiation and laser - Zhang 1991 sham laser acupuncture. Placebo methods may reduce participants' subjectivity but it is difficult to use, especially in China where many have previous experience of acupuncture and may well know the difference between real and pseudo-acupuncture (Li 2009; Yang 2009).

# 3. COMPARISON 1: ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Please see Summary of findings for the main comparison.

# 3.1 Global state

Although different criteria were used to rate 'not improved' all three to 12 month studies favoured the group into which acupuncture was added. The quality of data is not good but if, by this simple addition of acupuncture, even a proportion of the effect seen in these trials was real, that would be clinically important. This finding alone merits further testing.

# 3.2 Mental state

These trials presented 12 different ways of recording mental state. There were some exceptions and some heterogeneous results, however, essentially, the data favoured the acupuncture group. Most of the measures indicated a modest improvement derived from trials at moderate risk of bias, but findings that were statistically if not clinically significant. These data are certainly consistent with the 'global state' outcomes above and also merit further work.

# 3.3 Behaviour

Attrition was remarkably low. Such high rates of retention are unusual for trials involving people with schizophrenia and probably reflect the differences in approaches to care and the care culture, rather than the treatment interventions, however, the approach and care culture of China, in which these trials were conducted applies to at least 20% of the global population. Acupuncture trials of this sort are clearly are possible in China. More, larger and clearer studies are needed.

# 3.4 Service outcome

Only one of the 30 included studies reported this important outcome. Acupuncture added to standard dose antipsychotics did reduce average hospitalisation days compared with standard dose antipsychotics alone. Again, this is consistent with the above results and because it was a study with 120 participants of low quality attempts should be made to replicate this finding.

#### 3.5 Adverse effects

The wide-ranging Treatment Emergent Symptom Scale (TESS) scores were lower for those given acupuncture added to standard dose antipsychotic treatment. These results are based on a study

Acupuncture for schizophrenia (Review)

Copyright  ${\small ©}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.
size of only 90 participants (electro - Yao 2006), but, nevertheless, the general impression remains that acupuncture is not associated with noticeable adverse effects and also that it does little for any existing adverse effects of the medications.

### 4. COMPARISON 2: ACUPUNCTURE added to LOW DOSAGE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Please see Summary of findings 2. The low dose antipsychotic was chlorpromazine at less than 300 mg/day, risperidone 1 to 2 mg/d, clozapine 100 to 150 mg/d or, a reduction of about 60% of previously daily levels.

#### 4.1 Global state

Relapse, the primary outcome of this review, was only reported by acupoint inj - Pan 2002. Acupuncture added to low dose antipsychotics did reduce the number of participants experiencing relapse compared with standard dose antipsychotics (n = 170, 1 RCT, RR 0.57 CI 0.37 to 0.89). The study followed up relapse for two years but the result is questionable because of high risk of bias and incomplete outcome data. The outcome of 'no clinically important change' in global state did not show a benefit for acupuncture group and a similar result was found for the short-term CGI-SI. It is, therefore, difficult to be conclusive.

#### 4.2 Mental state

Rather weak and sometimes heterogeneous data suggest that the additional acupuncture did not really make much difference to participant's mental state. This impression remains even when we removed traditional - Wang 2006. the study that caused high heterogeneity.

#### 4.3 Behaviour

Again, very, very few people left these studies (~2%) indicating the different circumstances in which these trials were undertaken compared to the studies of Western medicine. Just because this indicates that the studies were different to that seen in the West does not mean the findings are inapplicable. Conversely, it may reflect that people did not feel empowered to leave, and that, perhaps with that dis-empowerment may come results of limited veracity.

### 4.4 Adverse effects

We found TESS endpoint score and RESES endpoint score were lower for people given acupuncture - but the RESES endpoint score was calculated according to the assumption of *The Cochrane Handbook or Systematic Reviews of Interventions* and could incorporate even more error than other data. Data for all other adverse effects are limited and not entirely convincing. It is not clear that the reduction of antipsychotic paired with acupuncture is helpful for a wide range of adverse effects.

### 5. COMPARISON 3: ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS

Please see Summary of findings 3.

### 5.1 Global state

The criteria for clinically important change of 'not improved' in global state was not standardised and findings were not consistent. The two trials using the same criteria found in favour of the acupuncture but the other three, employing different measures, were equivocal. This is surprising to those of us who are used to using standard dose antipsychotic drugs. Trials involving nearly 400 people indicate that acupuncture is as potent as antipsychotics - or more so. All studies are limited by their power and quality and highgrade replication is important.

### 5.2 Mental state

BPRS score at short-term based on two small studies, traditional -Wang 2006 and laser - Zhang 1991. Heterogeneity exists between traditional acupuncture and laser acupuncture subgroup and data from each study drew different conclusions. Findings tend to favour acupuncture for positive symptoms and neither group for negative. Again, if a real finding, these results are similar to those that are found for trials of new antipsychotic drugs. They do tend to be biased in favour of the experimental treatment, in this case acupuncture, but that does not mean that there is a real effect.

#### 5.3 Behaviour

Again, very high retention rates highlight the difference between these trials and what is seen in most of the rest of the world.

#### 5.4 Adverse effects

For this comparison adverse effect data were limited. As with the other comparisons there is no clear indication of adverse effects of acupuncture, although one study reported the laser acupuncture group experienced numbness over the ear, upper extremity and chest on the treated side.

## 6. COMPARISON 4: ACUPUNCTURE added to TCM DRUG versus TCM DRUG

Please see Summary of findings 4.

#### 6.1 Global state

When comparing traditional acupuncture plus capsule with capsule alone there was significant difference in global state (n = 360, 2 RCTs, RR 0.11 Cl 0.02 to 0.59) but the opposite result if comparing electroacupuncture plus decoction with decoction alone (n = 94, 1 RCT, RR 0.77 Cl 0.57 to 1.06). We have not really had the power to investigate the different types of acupuncture but this finding indicates that the traditional (Figure 1) may well have advantages over the electroacupuncture technique, also that the TCM drug may have little to offer, in itself, for the treatment of people with schizophrenia (Rathbone 2005b).

#### 6.2 Behaviour

Similar to the above comparisons - no participants left early. This is unusually high compliance compared to trials completed in the Western world.

#### 7. COMPARISON 5: ACUPUNCTURE versus TCM DRUG

Please see Summary of findings 5.

#### 7.1 Global state

Overall, there was no difference in the rates of not improved in global state, however, different approaches were used as well as different definitions of outcome. Two studies with the same criteria indicated that there was no significant difference between traditional acupuncture group and TCM drug group for one definition of 'not improved' (n = 360, 2 RCTs, RR 1.46 CI 0.74 to

Acupuncture for schizophrenia (Review)

2.87) but when another definition was employed a different result was found (worsening in global state, n = 88, 1 RCT, RR 0.52 CI 0.34 to 0.80, Analysis 5.1). The answer is clearly not stable and should be further investigated.

#### 7.2 Behaviour

No participants dropped out from either the acupuncture or TCM drug group. Low attrition probably refects, at least in part, that both acupuncture and Chinese herb medicine originate from China, have a long and trusted history, and these studies were undertaken in China.

### 8. COMPARISON 6: ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY

Please see Summary of findings 6.

#### 8.1 Behaviour

The only relevant study (n = 68) (EACT - Xue 1987), undertaken more than 20 years ago, reported no participants leaving within the 24 days of treatment. The short treatment period and a special approach to care may part of the reason for this.

#### 8.2 Adverse effects

Electric acupuncture convulsive therapy reduced spinal fracture rate but did not reduce back pain compared with electroconvulsive therapy. Both convulsive therapies were given without anaesthetic. We still can not draw firm conclusion from weak, and hopefully, outdated evidence.

#### 9. Missing outcomes

Even for Chinese medicine, which includes acupuncture, and has been used to treat schizophrenia-like illnesses for over 2000 years, we found no data relating to death, engagement with services, satisfaction with treatment, quality of life or economic outcomes (details in Table 4).

### Overall completeness and applicability of evidence

#### 1. Completeness

Thirty studies were included in this update review with more than 2500 participants. We compared acupuncture added to antipsychotic drugs (regular dosage or low dosage) with standard dose antipsychotics; acupuncture plus TCM drug with TCM drug alone; acupuncture with antipsychotic drugs or TCM drug; and also electric acupuncture convulsive therapy with electroconvulsive therapy. We found many outcomes were missing such as death, engagement with services, satisfaction with treatment, quality of life or economic outcomes. Most missing outcomes are participantoriented data.

Only one study reported our primary outcome, relapse, and even that was incomplete. One study reported a service outcome, time in hospital, which was not the pre-stated outcome of our review. This situation is unlikely to change until there is more wide agreement about the need for specific and clinically important outcomes.

#### 2. Applicability

The 30 included studies were published between 1986 and 2011 and almost exclusively completed in China where trial attrition is unusually low compared to the Western world. Also, although the great majority of people patients with schizophrenia are in the community most of included studies were undertaken in hospital settings from which attrition may be more difficult. Any findings of this review may be more applicable to people in hospital in China.

#### **Quality of the evidence**

Overall, the quality was not strong (Figure 3). Most studies did not adhere to the CONSORT statement; two-thirds of the included studies did not describe how they undertook the randomisation procedure and one sixth reported incomplete outcome data. Schizophrenia is a chronic illness but most studies reported shortterm data. We have tried to be generous in our judgements regarding quality and there is a risk that we, in doing this, have downplayed a real risk of bias in this group of studies and therefore in this review.

#### Potential biases in the review process

Please see above (Quality of the evidence). Otherwise, we used thorough search strategies and strictly followed the review protocol in the process of study selection, data extraction and analysis. We did employ only published reports and could not entirely avoid the potential for publishing bias for negative results and small studies.

Problems continue to exist for some of the included studies (acupoint inj - Pan 2002; electro - Zhang 1993). We found that data from Wu 2004 were extremely similar to those of acupoint inj - Pan 2002, with the exception of numbers in each group. The acupoint method and the BPRS's factor data were very similar and we were unable to gain further details about Wu 2004 and had to exclude that study in its entirety as a result.

# Agreements and disagreements with other studies or reviews

### 1. Previous version of this Cochrane review

This review updates and improves the previous version (Rathbone 2005a). We have changed the style of the Study ID tag to be more informative, using acupuncture's category plus author's surname plus publication date. The current version now includes 30 studies, whereas the previous version included only five.

The previous Cochrane review had considered two papers reporting two different studies; now electro - Zhang 1993 is the same as study 'Zhang 1994' from the previous version because we found an additional relevant reference (Zhang 1993). We changed 'Gang 1997' of the previous version to 'electro - Zhou 1997'.

#### 2. Other reviews

We identified two other acupuncture for schizophrenia reviews (Lee 2009; Nicholas 1997). The latter concluded that the results provided limited evidence overall and that the low quality studies did not allow any firm conclusions to be drawn and that international studies are needed to test whether there is any real effect. We agree with this view.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

## AUTHORS' CONCLUSIONS

### Implications for practice

### 1. For people with schizophrenia

Acupuncture has been used for treating schizophrenia-like illnesses for over 2000 years. Though acupuncture as a complementary treatment has become more popular in the West (Zhang 2012), 29 of the 30 included studies were undertaken in China and the findings are more appropriate to a Chinese population. For this update, new studies were found to compare acupuncture added to a Traditional Chinese Medicine (TCM) drug with TCM drug, acupuncture with TCM drug, electric acupuncture convulsive therapy with electroconvulsive therapy and we separately analysed acupuncture added to standard dose antipsychotics versus standard dose antipsychotics or acupuncture adde to low dose antipsychotics versus standard dose antipsychotics. The limited evidence suggests that combining acupuncture with standard dose antipsychotics might benefit the individual's mental state especially in the short-term. Acupuncture appears to have some benefit related to relapse when added to low dose antipsychotics compared to standard dose antipsychotics along with a reduction of drug-related adverse effects.

### 2. For clinicians

Adopting acupuncture to treat schizophrenia needs co-operation between psychiatrists and acupuncturists. Few data were available making the comparisons between acupuncture and antipsychotics, between acupuncture added to TCM drug and TCM, between acupuncture and TCM drug, and between electric acupuncture convulsive therapy and electroconvulsive therapy limited. Acupuncture treatment effects consists of three effects special effects, non-special effects and psychological effects (Yang 2012). Without enough evidence, we could not draw conclusions if the effect of acupuncture when combining with antipsychotics is a placebo response. Without enough evidence we remain confused about the interaction between acupuncture and low dosage antipsychotic medication. Placebo and recognised effective drug-control in three-arm trials need to be conducted to provide strong evidence.

It is surprising that most of the studies used the Treatment Emergent Symptom Scale (TESS) or Rating Scale for Extrapyramidal Side Effects (RESES) to assess adverse effects, but few focused on acupuncture's relative adverse effects such as skin allergies. Though acupuncture has a long history, this intervention is still without a special acupuncture-relative adverse effects scale. Adverse effects of acupuncture for schizophrenia are inadequate.

### 3. For managers of policy makers

Acupuncture as a valuable complementary medicine is gradually affecting the global medical treatment system. Only one study reported service outcomes and indicated that using acupuncture added to standard dose antipsychotics shorten hospitalisation days than using standard dose antipsychotics alone. This traditional treatment originating from ancient Chinese culture needs more real-word trials to evaluate the treatment method objectively and completely, not limited by assessing treatment effects and adverse effects but also focusing on social and economic conditions.

### Implications for research

### 1. General

All studies should now comply with the Consolidated Standards of Reporting Trials (CONSORT, Moher 2001). More transparency in the reporting of randomised controlled trials, would enable readers to understand the design, conduct, analysis and interpretation, and to assess the validity of results. Although binary data is easier to interpret, where continuous data are used, some measure of variance should be provided. Data presented in graphs should be accompanied by exact numbers and standard deviations in the text.

### 2. Specific

### 1. Reviews

There are further refinements of this review that could take place with better inclusion of additional data (Table 1, Table 2) and possibility of additional comparisons (Table 5). This review is already large and, in the future, it may be best to split this work into separate reviews per comparison with an 'overview' review.

### 2. Trials

In 2010, the STRICTA Group, CONSORT Group and the Chinese Cochrane Centre collaborated to develop STRICA (MacPherson 2010) to ensure that acupuncture trials are more accurately interpreted and more easily replicated. Further research is essential to inform both clinicians and patients about the effects of acupuncture in the treatment of schizophrenia. Trial methodology should follow both CONSORT (Moher 2001) and STRICTA guidelines.

Acupuncture is an important intervention that is likely to be widely used, at least, in China. Considering the limited data in this review, we do think that further large simple trials are indicated. This update highlights the need for standardised clinical trials, which evaluate acupuncture for people with schizophrenia, which thoroughly investigate effects of different categories of acupuncture, which investigate interaction in combination regimens, with trialists employing appropriate realworld methodology. We suggest such a design in Table 6.

There are, however, specific issues to consider.

### 2.1 Standardisation of acupuncture

We included 30 studies and five categories of acupuncture according to different stimulation methods and one according to special acupuncture theory application method. The majority used different acupoints and practice procedures. Although acupuncture is an important part of TCM and has the typical characteristics of this approach - individualised treatment - to objectively evaluate the effects of acupuncture standardisation of technique would be useful.

### 2.2 Blinding method and sham acupuncture

Human behaviour is influenced by what we know or believe and it is a particular risk of human healthcare research. The use of blinding methods makes it more difficult to bias results intentionally or unintentionally and helps ensure credibility of study conclusions (Simon 2000). Eight studies adapted a blinding method but only four used sham acupuncture. Though appropriate sham acupuncture is a key factor for acupuncture trials, it can also be a barrier for the trialist (Li 2009). More than 20 studies

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



in this review probably have some degree of performance bias and only seven used blinded assessors. As acupuncture is an externally applied treatment to which special theory is associated, patient-blinding (sham acupuncture) may be difficult, but assessorblinding is not so problematic. At the very least, to minimise subjectivity, assessor-blinding does need to be practiced in further clinical trials.

### 2.3 Control group

A placebo-controlled (appropriate sham acupuncture) trial is beneficial to confirm absolute effects of acupuncture. We did not find any study of this comparison. In this version of review we reported results of:

- acupuncture added to standard dose antipsychotics versus standard dose antipsychotics;
- acupuncture added to low dose antipsychotics versus standard dose antipsychotics;
- acupuncture versus standard dose antipsychotics;
- acupuncture added to TCM drug versus TCM drug;
- acupuncture versus TCM drug; and a special comparison
- electric acupuncture convulsive therapy versus electroconvulsive therapy.

Some of the control group treatments are definitely effective (such as antipsychotics; Rattehalli 2010; Adams 2007), but others may be more like a placebo group (e.g. TCM drug - Rathbone 2005b). To evaluate the effects of acupuncture choosing recognised valid treatment as the control group would be beneficial to obtain highquality globally applicable evidence.

### ACKNOWLEDGEMENTS

This review was initiated by John Rathbone who wrote the protocol, selected studies, data extracted, assimilated data and wrote the final first report (Rathbone 2005a). Jun Xia helped select studies, data extract, translate text and contact for that first version.

Many thanks to Samantha Roberts for the 2012 trials search. We are most grateful for the ongoing support of Claire Irving at the Cochrane Schizophrenia Group's editorial base.

The Cochrane Schizophrenia group provide a standard template for their methods section and other parts of the protocol. We have used this template and adapted it accordingly.

Authors Yuyin Pan, Cheng Luo, Guimei Cui, Jing Cheng and Professor Xueguang Bai were kind enough to respond to our telephone or e-mail inquiries, for which we are very grateful.

### REFERENCES

#### References to studies included in this review

#### acupoint cat - Sun 2005 {published data only}

Sun L, Zhang H, Xia S, Gong F, Wang J, Guo M, et al. Control study of hallucinations of schizophrenia by the acupoint catgut treatment [穴位埋线治疗分裂症幻觉的对照研究]. *Practical Clinical Journal of Integrated Traditional Chinese and Western Medicine* [#########] 2005;**5**(6):5-6.

#### acupoint cat - Wang 1997 {published data only}

Wang J, Ye Y. Effective analysis of Tinggong embedding catgut method combining with antipsychotics on auditory hallucinations treatment [听宫穴埋线法合并抗精神病 药治疗幻听的疗效分析]. Chinese Journal of Psychiatry 1997;**30**(4):202.

#### acupoint inj - Pan 2002 {published data only}

\* Pan Y, Wang J, Liu S, Wu C, Wang L, Shen L. Treatment for schizophrenia patients with acupoint injection [穴位注药治疗精神分裂症的研究]. Journal of Taishan Medical College 2002;23(2):128-30.

Pan Y, Wu C, Pan Y. A control study of schizophrenia patients according to types [精神分裂症分型施治的对照研究]. Journal of the Handan Medical College 2003;**16**(6):493-4.

Pan Y, Pan Y, Wu C, Li M. A follow-up study of schizophrenia patients treated by acupuncture and drug [针药并用治疗精神分裂症的随访研究]. *Health Psychology Journal* 2002;**10**(6):456-7.

#### acupoint inj - Wang 2000 {published data only}

Wang G, Ji R, Pei G. The control study of injection of clonazepami into T'ingkung for refractory auditory hallucination of schizophrenia [听宫穴注射氯硝西泮治疗精神分裂症顽固性幻听对照研究]. Journal of Clinical Psychosomatic Diseases [########] 2000;6(3):143-4.

#### acupoint inj - Yang 2000 {published data only}

Yang S, Liu G. Observation on intractable auditory hallucination treated by injecting sulpiride into acupoints [穴位注射舒必利治疗顽固性幻听疗效观察]. Journal of Practical Traditional Chinese Medicine 2000;**16**(7):24-5.

#### EACT - Xue 1987 {published data only}

Xue C, Wang J, Tang M. Comparative study of spinal fractures in electric acupuncture convulsive therapy, electroconvulsive therapy and epilepsy [电针抽搐治疗、电抽搐治疗与癫 痫所致的脊柱骨折]. *Chinese Journal of Neurology and Psychiatry [Chung Hua Shen Ching Ching Shen Ko Tsa Chih]* 1987;**20**(6):346-9.

#### electro - Chen 2006 {published data only}

Chen Z, Song H, Wen N. [Title only available in Chinese characters] [电针治疗精神分裂症后抑郁]. Modern Journal of Integrated Traditional Chinese and Western Medicine 2006;**15**(13):1776-7.

#### electro - Chen 2008 {published data only}

Chen K, Li D, Ye L. The effect of electric acupuncture combined with aripiprazole in treating negative symptoms of schizophrenia [电针合并阿立哌唑治疗精神分裂症阴性症状的 疗效观察]. *Medical Journal of Chinese People's Health* [######] 2008;**20**(1):7-10.

#### electro - Cheng 2009 {published data only}

Bai X. An 8-week randomized study of acupuncture in the treatment of auditory hallucinations in refractory and non-refractory schizophrenic patients. Stanley Foundation Research Programs. China, 2002.

\* Cheng J, Wang G, Xiao L, Wang H, Wang X, Li C. Electroacupuncture versus sham electro-acupuncture for auditory hallucinations in patients with schizophrenia: a randomized controlled trial. *Clinical Rehabilitation* 2009;**23**(7):579-88.

#### electro - Cui 2000 {published data only}

Cui G, Wang P. The clinical study of combining electrostimulation and chlorpromazine in schizophrenia [电针合并氯丙嗪治疗精神分裂症的临床研究]. *Modern Journal of Integrated Chinese Traditional and Western Medicine* 2000;**9**(16):1536-7.

#### electro - Ding 2005 {published data only}

Ding G, Ling F, Zhang J. [only Chinese Title available] [针灸 治疗慢性精神分裂症的临床效果分析]. Modern Journal of Integrated Traditional Chinese and Western Medicine 2005;**14**(1):53-4.

#### electro - Wang 2005 {published data only}

Wang P, Wang X, Li X, Wu X, Duan D. Controlled studies on combination of electroacupuncture and risperidone in negative symptoms of schizophrenia [电针合并利培酮治疗精神分裂 症阴性症状对照研究]. *Liaoning Journal of Traditional Chinese Medicine* 2005;**32**(7):710-1.

#### electro - Xiong 2010 {published data only}

Xiong D, Liu L, Yi Y, Ye F. Observation on the therapeutic effect of electroacupuncture combined with small dose of clozapine in clinical treatment of refractory schizophrenia [电针联合小剂量氯氮平治疗难治性精神分裂症的疗效观察]. Chen Tzu Yen Chiu / Acupuncture Research 2010;**35**(2):134-7. [MEDLINE: 20626147]

#### electro - Yao 2006 {published data only}

Yao F, Sun F, Zhang Z. Short-term curative effect of electroacupuncture as an adjunctive treatment on schizophrenia [电针辅助治疗精神分裂症的近期疗效观察]. *Chinese Journal of Integrated Traditional and Western Medicine* 2006;**26**(3):253-5. [MEDLINE: 16613275]

#### electro - Zhang 1987 {published data only}

Wu T. Personal Communication 2008.

\* Zhang L, Tang Y, Zhu W, Xu S. Comparative study of schizophrenia treatment with electroacupuncture, herbs and chlorpromazine. *Chinese Medical Journal* 1987;**100**:152-7.

#### Acupuncture for schizophrenia (Review)



Zhang L, Xu S, Tang Y, Zhu W. A comparative study of the treatment of schizophrenia with electric acupuncture, herbal decoction and chlorpromazine. *American Journal of Acupuncture* 1990;**18**(1):11-4. [MEDLINE: 99343121; PMID 10416732]

### electro - Zhang 1993 {published data only}

\* Zhang B, Meng S, Yu J, Quan C, Li W, Sun H, et al. A controlled study of therapeutic effects of computer-controlled electric acupuncture treatment on refractory schizophrenia. *World Journal of Acupuncture-Moxibustion* 1993;**3**(4):3-9.

Zhang B, Meng S, Yu J, Quan C, Li W, Sun H, et al. Clinical therapeutic effect of mentality electroacupuncture on schizophrenia [智能电针仪治疗精神分裂症临床疗效对照研究]. *Chinese Acupuncture and Moxibustion* 1994;**17**(1):17-20. [MEDI9402]

#### electro - Zhang 2001 {published data only}

Zhang Y, Shao H, Zhao X, Liu W, He J, Chai L, et al. The clinical efficacy with intelligent electro-acupuncture of treating schizophrenia with depressive symptoms [智能电针治疗精神分裂症伴发抑郁症状的临床疗效观察]. Chinese Journal of Behavioral Medical Science 2001;**10**(1):44-5. [MEDI0104]

#### electro - Zhou 1997 {published data only}

Zhou G, Jin S, Zhang L. Comparative clinical study on the treatment of schizophrenia with electroacupuncture and reduced doses of antipsychotic drugs. *American Journal of Acupuncture* 1997;**25**(1):25-31. [153rd Annual Meeting of the American Psychiatric Association [CD-ROM]: MARATHON Multimedia, 2000 NR80]

#### laser - Liu 1986 {published data only}

Liu Z, Wang Y, Zhang S, He A, Chen Y, Liu X. Therapeutic effect of He-Ne laser irradiation of point erman in schizophrenic auditory hallucination--a clinical assessment. *Journal of Traditional Chinese Medicine = Chung i tsa chih ying wen pan / sponsored by All-China Association of Traditional Chinese Medicine, Academy of Traditional Chinese Medicine* 1986;**6**(4):253-6. [PUBMED: 3600018]

#### laser - Ma 1999 {published data only}

Ma Z, Li X, Lv Y. A control study of auditory hallucination treated with point-stimulating therapy of helium neon [氦氖激光穴 位照射治疗具有幻听症状的精神分裂症对照研究]. Sichuan Mental Health [######] 1999;**12**(2):90-1.

#### laser - Zhang 1991 {published data only}

Zhang B, Quan C, Yu J, Li W, Sun H, Liu S, et al. A controlled study of clinical therapeutic effects of laser acupuncture for schizophrenia [激光针灸治疗精神分裂症临床疗效的对照 研究]. *Chung Hua Shen Ching Ching Shen Ko Tsa Chi (Chinese Journal of Neurology and Psychiatry)* 1991;**24**(2):81-3, 124. [MEDLINE: 91317070; PMID 1860386]

### traditional - Bouhlel 2011 {published data only}

Bouhlel S, El-Hechmi S, Ghanmi L, Ghaouar M, Besbes C, Khaled M, et al. Effectiveness of acupuncture in treating schizophrenia: A clinical randomised trial about 31 patients. *Tunisie Medicale* 2011;**89**(10):774-8.

#### traditional - Liu 2010 {published data only}

Liu X. Clinical observation of acupuncture treatment in 50 schizophrenia with refractory auditory hallucinations [针刺 治疗精神分裂症顽固性幻听患者50例临床观察]. Shanghai Journal of Traditional Chinese Medicine [Shang Hai Zhong Yi Yao Za Zhi][####] 2010;**7**:621-4.

#### traditional - Luo 2006 {published data only}

Luo C, Zhou W. Study of acupuncture adjunctive therapy on type II syndrome of schizophrenia [针灸辅助治疗精神分裂症 II型综合征的研究]. *Modern Journal of Integrated Traditional Chinese and Western Medicine* 2006;**15**(2):148-9.

#### traditional - Ma 2008 {published data only}

Ma L, Zhang E, Lu S. Control study of acupuncture combining with risperidone for schizophorenia [针刺合用利培酮治疗精神 分裂症对照研究]. *Medical Journal of Chinese People's Health* [# #####] 2008;**20**(17):1981.

#### traditional - Tang 2005 {published data only}

\* Tang Y, Huang Q, Guo J. Clinical analysis of acupuncture for auditory hallucinations [针刺治疗幻听临床分析]. *Chinese Journal of Current Traditional and Western Medicine* 2005;**3**(5):431-2.

Tang Y, Huang Q, Li H, Wen Y, Guo J. Clinical analysis of acupuncture in the treatment of phonism [针刺治疗幻听临 床分析]. *Chinese Journal of Health Psychology [########]* 2007;**15**(4):367.

#### traditional - Wang 2006 {published data only}

Wang L, Xie Y. Clinical study of acupuncture for hebephrenic schizophrenia [针刺治疗青春期精神分裂症的临床研究]. Journal of Clinical Acupuncture and Moxibustion 2006;**22**(9):12-4.

#### traditional - Xu 2004 {published data only}

Xu T, Diao H, Xu P, Gan H, Sun Z, Xu L, et al. Clinical study on acupuncture combined with small dose of antipsychotics for treatment of 40 cases of schizophrenia [針刺配合小劑量 抗精神病藥物治療精神分裂癥40例臨床研究]. Journal of Traditional Chinese Medicine [Chung i tsa chih ying wen pan][### #] 2004;**45**(1):22-5.

#### traditional - Zhao 2005a {published data only}

Zhao Y. Acupuncture Is It Recover Health Capsule Trear Schizophrenia and Influence to Patient's Blood Free Radical Supersession to Share [针刺合用复元康胶囊治疗精神分裂 症及其对患者血自由基代谢影响]. Doctoral Dissertation of Heilongjiang University of TCM 2005.

#### traditional - Zhao 2005b {published data only}

Zhao Y. Acupuncture Is It Recover Health Capsule Trear Schizophrenia and Influence to Patient's Blood Free Radical Supersession to Share [针刺合用复元康胶囊治疗精神分裂 症及其对患者血自由基代谢影响]. Doctoral Dissertation of Heilongjiang University of TCM 2005.

Acupuncture for schizophrenia (Review)



### References to studies excluded from this review

#### Luo 2006a {published data only}

Luo C, Zhang Y, Xiong K, Zhou W. A study of acupunncture on prevent side - effect caused by antipsychotics [针刺预防抗精神 病药物副反应110例临床观察]. Journal of the Yunnan College of Traditional Chinese Medicine 2006;29(3):27-8.

#### Ma 2002 {published data only}

Ma H, Guo Y, Fan H, Li Y, Yue S. [Title only available in Chinese characters] [头部电针治疗慢性精神分裂症临床研究]. Proceedings of the 7th Psychiatry Conference of the Society of Intergreted Traditional Chinese and Western Medicine. 2002:213-6.

Yang S, Guo Y, Yue S. [Title only available in Chinese characters] [头部电针治疗慢性精神分裂症临床疗效及护理]. Medical Journal of Chinese People's Health [######] 2003;**15**(10):630-1.

#### Ma 2004 {published data only}

Ma L. [Title only available in Chinese characters] [电针百会、 神庭穴对精神分裂症患者认知功能影响的临床研究]. MSc dissertation submitted to the Beijing University of Chinese Medicine, China 2004.

\* Ma L, Gu S, Zhang X. Effect of electroacupuncture at baihui and shenting on cognitive function of patients with convalescent schizophrenia. *Chinese Journal of Clinical Rehabilitation* 2005;**9**(4):99-101.

### Sun 1994 {published data only}

Sun Z, Wu H, Sun Y, Zhao A. Observation on treatment of 350 cases of schizophrenia with electroacupuncture and acupointinjection. Chinese Acupuncture and Moxibustion 1994, issue S1:53-4.

#### Wu 2004 {published data only}

Wu C, Li M. A study of acupuncture point injection in the treatment of schizophrenia. *Chinese General Practice* [######] 2004;**7**(19):1382-4.

#### Xiong 2009 {published data only}

Xiong D, Yi Y, Zhu X, Hu W. Controlled study on therapeutic effects of electroacupuncture and modified electroconvulsive therapy on catatonic schizophrenia [电针与无抽搐电休克治疗紧张型精神分裂症]. Chinese Acupuncture and Moxibustion 2009;**29**(10):804-6. [MEDLINE: 19873916]

#### Xue 1985 {published data only}

Xue C, Xie H, Ruan Q, Cheng Y, Liu D. Electric acupuncture convulsive therapy. *Convulsive Therapy* 1985;**1**(4):242-51. [MEDLINE: 94072826; PMID 8251722]

#### Zhong 1995 {published data only}

Zhong H, Feng X, Luo H, Jia Y, Zhao X. Comparative observation on treatment of psychosis with electrode and electroacupuncture controlled by processors. *World Journal Acupuncture-Moxibustion* 1995;**5**(3):24-7. [MEDI95S5]

#### Zhuge 1993 {published data only}

Zhuge D, Chen J. Comparison between electro-acupuncture with chlorpromazine and chlorpromazine alone in 60

Cochrane Database of Systematic Reviews

schizophrenic patients. *Chung-Kuo Chung Hsi i Chieh Ho Tsa Chih* 1993;**13**(7):408-9. [MEDLINE: 94072826; PMID 8251722]

#### **References to studies awaiting assessment**

#### NCT01167348 {published data only}

NCT01167348. The effectiveness of auricular acupressure on bodybweight parameters on patients with schizophrenia. http://ClinicalTrials.gov/show/NCT01167348 2010.

#### Additional references

### Adams 2007

Adams EC, Awad G, Rathbone J, Thornley B. Chlorpromazine versus placebo for schizophrenia. *Cochrane Database of Systematic Reviews* 2007, Issue 2. [DOI: 10.1002/14651858.CD000284]

#### Alderson 2004

Alderson P, Green S, Higgins JPT. Cochrane Reviewers' Handbook 4.2.2 [updated December 2003]. The Cochrane Library. Chichester, UK: John Wiley & Sons, Ltd, 2004. [In: The Cochrane Library, Issue 1, 2004. Chichester, UK: John Wiley & Sons, Ltd.]

#### Altman 1996

Altman DG, Bland JM. Detecing skewness from summary information. *BMJ* 1996;**313**:1200. [OLZ020600]

#### Andreasen 1982

Andreasen NC. Negative symptoms in schizophrenia. Definition and reliability. *Archives of General Psychiatry* 1982;**39**:784-8.

#### Bland 1997

Bland JM. Statistics notes. Trials randomised in clusters. *BMJ* 1997;**315**:600.

#### Boissel 1999

Boissel JP, Cucherat M, Li W, Chatellier G, Gueyffier F, Buyse M, et al. The problem of therapeutic efficacy indices. 3. Comparison of the indices and their use [Apercu sur la problematique des indices d'efficacite therapeutique, 3: comparaison des indices et utilisation. Groupe d'Etude des Indices D'efficacite]. *Therapie* 1999;**54**(4):405-11. [PUBMED: 10667106]

### Chen 2002

Chen Y. Chines Classification of Mental Disorders (CCMD-3): Towards Integration in International Classification. *Psychopathology* 2002;**35**(2-3):171-5.

#### Deeks 2000

Deeks J. Issues in the selection for meta-analyses of binary data. Proceedings of the 8th International Cochrane Colloquium; 2000 Oct 25-28; Cape Town. Cape Town: The Cochrane Collaboration, 2000.

Acupuncture for schizophrenia (Review)



#### Divine 1992

Divine GW, Brown JT, Frazier LM. The unit of analysis error in studies about physicians' patient care behavior. *Journal of General Internal Medicine* 1992;**7**(6):623-9.

#### Donner 2002

Donner A, Klar N. Issues in the meta-analysis of cluster randomized trials. *Statistics in Medicine* 2002;**21**:2971-80.

#### Egger 1997

Egger M, Davey-Smith G, Schneider M, Minder CSO. Bias in meta-analysis detected by a simple, graphical test. *BMJ* 1997;**13**:629-34.

#### Elbourne 2002

Elbourne D, Altman DG, Higgins JPT, Curtina F, Worthingtond HV, Vaile A. Meta-analyses involving crossover trials: methodological issues. *International Journal of Epidemiology* 2002;**31**(1):140-9.

#### Ernst 2001

Ernst E, A White. Prospective studies of the safety of acupuncture: a systematic review. *American Journal of Medicine* 2001;**110**:481-5.

#### Fan 2010

Fan F, Hong X, Song J. Research and prospect on laser acupuncture [激光针灸的研究现状与展望]. Laser Journal 2010;**31**(5):58-9.

#### Furukawa 2006

Furukawa TA, Barbui C, Cipriani A, Brambilla P, Watanabe N. Imputing missing standard deviations in meta-analyses can provide accurate results. *Journal of Clinical Epidemiology* 2006;**59**(7):7-10.

### Gulliford 1999

Gulliford MC. Components of variance and intraclass correlations for the design of community-based surveys and intervention studies: data from the Health Survey for England 1994. *American Journal of Epidemiology* 1999;**149**:876-83.

#### Guo 2010

Guo Y, Shi Y, Li Y, Liu Y, Zhang Y. Research progress of Traditional Chinese Medicine on schizophrenia treatment [中药治疗精神 分裂症的研究进展]. *Modern Journal of Integrated Traditional Chinese and Western Medicine* 2010;**19**(27):3543-5.

#### Guy 1970

Guy W, Bonato RR, eds. Clinical global impressions. Manual for the ECDEU Assessment Battery 2. Rev ed. National Institute of Mental Health, 1970.

#### Guy 1976

Guy W. ECDEU Assessment Manual for Psychopharmacology (DOTES: Dosage Record and Treatment Emergent Symptom Scale). Rockville: National Institute of Mental Health, 1976.

#### Haddock 1999

Haddock G, McCarron J, Tarrier N, Faragher EB. Scale to measure dimensions of hallucinations and delusions: the

psychotic symptom rating scales (PSYRATS). *Psychological Medicine* 1999;**29**(4):879-89.

#### Hamilton 1967

Hamilton M. Development of a rating scale for primary depressive illness. *British Journal of Social and Clinical Psychology* 1967;**4**(Dec, 6):278-96.

#### **Higgins 2003**

Higgins JP, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. *BMJ* 2003;**327**:557-60.

### Higgins 2011

Higgins JPT, Green S (editors). Cochrane Handbook for Systematic Reviews of Interventions Version 5.0.2 [updated September 2011]. The Cochrane Collaboration, 2011. Available from www.cochrane-handbook.org.

### Hutton 2009

Hutton Jane L. Number needed to treat and number needed to harm are not the best way to report and assess the results of randomised clinical trials. *British Journal of Haematology* 2009;**146**(1):27-30.

#### Kaptchuk 2002

Kaptchuk TJ. Acupuncture: theory, efficacy, and practice. American College of Physicians - American Society of Internal Medicine 2002;**136**:374-83.

#### Kay 1986

Kay SR, Opler LA, Fiszbein A. Positive and Negative Syndrome Scale (PANSS) Manual. North Tonawanda, NY: Multi-Health Systems, 1986.

#### Lee 2001

Lee S. From diversity to unit. The classification of mental disorders in 21st-century China. *Psychiatric Clinics of North America* 2001;**24**(3):421-31.

#### Lee 2009

Lee MS, Shin BC, Ronan P, Ernst E. Acupuncture for schizophrenia: a systematic review and meta-analysis. *International Journal of Clinical Practice* 2009;**63**(11):1622-33. [PUBMED: 19832819]

#### Leucht 2005

Leucht S, Kane JM, Kissling W, Hamann J, Etschel E, Engel RR. What does the PANSS mean?. *Schizophrenia Research* 2005;**79**(2-3):231-8. [PUBMED: 15982856]

### Leucht 2005a

Leucht S, Kane JM, Kissling W, Hamann J, Etschel E, Engel R. Clinical implications of brief psychiatric rating scale scores. *British Journal of Psychiatry* 2005;**187**:366-71. [PUBMED: 16199797]

#### Leucht 2007

Leucht S, Engel RR, Bauml J, Davis JM. Is the superior efficacy of new generation antipsychotics an artifact of LOCF?. *Schizophrenia Bulletin* 2007;**33**(1):183-91. [PUBMED: 16905632]

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



#### Li 2009

Li J. Research progress of placebo method in acupuncture clinical trials [安慰方法在针灸临床试验中的研究进展]. Journal of Complementary and Alternative Medicine 2009;**25**(1):53-4.

### Lu 2006

Lu W, Shi J, Zhou X, Chen F. Effective observation of body acupuncture combine with auricular acupuncture to treat schizophrenia refractory auditory hallucinations [体针合并 耳穴压籽治疗精神分裂症顽固性幻听的疗效观察]. Sichuan Mental Health 2006;**19**(4):236-7.

### MacPherson 2001

MacPherson H, Thomas K, Walters S, Fitter M. The York acupuncture safety study: prospective survey of 34 000 treatments by traditional acupuncturists. *2001* BMJ;**323**(Sept):486-7.

#### MacPherson 2010

MacPherson H, Altman DG, Hammerschlag R, Youping L, Taixiang W, White A, et al. STRICTA Revision Group. Revised STandards for Reporting Interventions in Clinical Trials of Acupuncture (STRICTA): extending the CONSORT statement. *Acupuncture in Medicine* 2010;**28**:83-93.

### Marshall 2000

Marshall M, Lockwood A, Bradley C, Adams C, Joy C, Fenton M. Unpublished rating scales: a major source of bias in randomised controlled trials of treatments for schizophrenia. *British Journal of Psychiatry* 2000;**176**:249-52.

#### Meng 2012

Meng S, Lv J, Wen X. Recent state on clinical application of acupoint catgut treatment [穴位埋线疗法临床应用近况]. Shanxi Journal of Traditional Chinese Medicine 2012;28(2):56-8.

#### Ming 2001

Translated by Zhu Ming. The Medical Classics of the Yellow Emperor. Beijing: Foreign Language Press, 2001.

#### Moher 2001

Moher D, Schulz KF, Altman D. The CONSORT statement: revised recommendations for improving the quality of reports of parallel-group randomized trials. *JAMA* 2001;**285**:1987-91.

### Nicholas 1997

Beecroft N, Rampes H. Review of acupuncture for schizophrenia. *Acupuncture in Medicine* 1997;**15**:91-4.

#### Overall 1962

Overall JE, Gorham DR. The Brief Psychiatric Rating Scale. *Psychological Reports* 1962;**10**:799-812.

#### Rathbone 2005b

Rathbone J, Zhang L, Zhang M, Xia J, Liu X, Yang Y. Chinese herbal medicine for schizophrenia. *Cochrane Database of Systematic Reviews* 2005, Issue 4. [DOI: 10.1002/14651858.CD003444]

#### Rattehalli 2010

Rattehalli RD, Jayaram MB, Smith M. Risperidone versus placebo for schizophrenia. *Cochrane Database of Systematic Reviews* 2010, Issue 1. [DOI: 10.1002/14651858.CD006918.pub2]

#### Schünemann 2008

Schünemann HJ, Oxman AD, Vist GE, Higgins JPT, Deeks JJ, Glasziou P, et al. Chapter 12: Interpreting results and drawing conclusions. In: Higgins JPT, Green S editor(s). Cochrane Handbook for Systematic Reviews of Interventions. The Cochrane Collaboration, 2008:359-83.

#### Shi 2007

Shi X. Science of Acupuncture and Moxibustion. Beijing: Chinese Publishing House of Traditional Chinese Medicine, 2007.

#### Shi 2010

Shi J, Zhou X, Gao M, Wang P. Status of acupuncture for schizophrenia [针灸治疗精神分裂症的现状]. *Medical Journal of National Defending Forces in North China* 2010;**22**(4):100-2.

#### Simon 2000

Simon JD, Douglas GA. Blinding in clinical trials and other studies. *BMJ* 2000;**321**:19-26.

#### Simpson 1970

Simpson GM, Angus JWS. A rating scale for extrapyramidal side effects. *Acta Psychoiatrica Scandinavica* 1970;**212**:11-9.

#### Tang 2010

Tang Z, Chen H. Research progress of effects and mechanism of acupoint injection [穴位注射作用效应及机制的研究进展]. *Zhejiang JITCWM* 2010;**20**(2):119-20.

#### **The Psychosis Professional Committee 1988**

The Psychosis Professional Committee of Chinese Integrative Medicine Association. The standard of integrative medicine syndrome type of schizophrenia [精神分裂症的中西医结 合辨证分型标准]. Chinese Journal of Integrative Medicine 1988;8(2):127.

#### Ukoumunne 1999

Ukoumunne OC, Gulliford MC, Chinn S, Sterne JAC, Burney PGJ. Methods for evaluating area-wide and organistation-based intervention in health and health care: a systematic review. *Health Technology Assessment* 1999;**3**(5):1-75.

### Wang 2007

Wang Y. Overview of Chinese Medicine and acupuncture for schizophrenia [精神分裂症的中药及针灸治疗概况]. Yunnan Journal of Traditional Chinese Medicine and Materia Medica 2007;**28**(10):54-5.

### Xia 2009

Xia J, Adams CE, Bhagat N, Bhagat V, Bhoopathi P, El-Sayeh H, et al. Loss to outcomes stakeholder survey: the LOSS study. *Psychiatric Bulletin* 2009;**33**(7):254-7.

### Xu 1991

Xu X. The English-Chinese Encyclopedia of Practical Traditional Chinese Medicine. Beijing: Higher Education Press, 1991.

Acupuncture for schizophrenia (Review)



#### Xu 2010

Xu T, Su J. Clinical status and thinking of acupuncture for schizophrenia [针灸治疗精神分裂症的临床现状与思考]. *CJITWM* 2010;**30**(11):1130-2.

#### Yang 2009

Yang Z, Zheng Q, Li Y, Wu X. Overview of adoption and research of sham acupuncture in acupuncture clinical research [针灸临 床研究中假针刺的运用及研究概况]. Journal of New Chinese Medicine 2009;41(3):93-5.

#### Yang 2012

Yang Y, Wei Y, Wang Y, Yin L, Xu Y, Liu Y. Current status and fundamental strategies for clinical research on acupuncture and moxibustion [针灸临床研究现状与基本策略]. Shanghai Journal of Traditional Chinese Medicine 2012;46(1):7-10.

#### Zhang 1993

Zhang B, Meng S, Yu J, Quan C, Li W, Sun H, et al. A controlled study of therapeutic effects of computer-controlled electric acupuncture treatment on refractory schizophrenia. World Journal of Acupuncture-Moxibustion 1993;3(4):3-9.

### CHARACTERISTICS OF STUDIES

Characteristics of included studies [ordered by study ID]

Zhang	1996
-------	------

Zhang J. Current status of integrated traditional and Western medicine study on schizophrenia. Chinese Journal of Integrated Traditional and Western Medicine 1996;16(11):643-5.

### Zhang 2012

Zhang Y, Lao L, Ceballos R. Acupuncture use among American adults: What acupuncture practitioners can learn from National Health Interview Survey 2007?. Evidence-Based Complementary and Alternative Medicine 2012. [DOI: 10.1155/2012/710750]

#### Zung 1965

Zung WW. A self-rating depression scale. Archives of General Psychiatry 1965;12:63-70.

#### References to other published versions of this review

#### Rathbone 2005a

Rathbone J, Xia J. Acupuncture for schizophrenia. Cochrane Database of Systematic Reviews 2005, Issue 4. [DOI: 10.1002/14651858.CD005475]

\* Indicates the major publication for the study

acupoint cat - Sun 200	5
Methods	Allocation: randomised. Blindness: not reported. Duration: 6 weeks.
Participants	Diagnosis: schizophrenia with auditory hallucinations (CCMD-3). N = 180. Age: mean age (28.10 ± 6.90) years (catgut treatment + low dose risperidone group); (30.40 ± 10.72) years (risperidone group). Sex: 86 women and 94 men.
	History: average duration of illness (5.85 ± 4.80) months (acupoint catgut treatment + low dose risperi- done group); (4.92 ± 3.69) years (risperidone group). Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-3 diagnosis of schizophrenia; the duration of illness more than 3 months and auditory hallucinations continued longer than 1 month; never received any antipsychotics or washout period was longer than 1 week; age between 18 and 60 years.
	Exclusion criteria: elderly and frail; pregnant women; severe somatic diseases; cerebral organic dis- eases.
Interventions	1. Acupoint catgut treatment + low dose risperidone: acupoint catgut treatment (Tinggong; opened mouth; once 7-10 days; 6 weeks as a treatment course) + low dose risperidone (1 to 2 mg/d; average dose (1.38 ± 0.52) mg/d). N = 88.
	2. Risperidone: initial dose 1 mg, adjusted the dose according to treatment courses and side effects and achieved to 4 to 10 mg/d within 2 weeks; average dose (5.95 ± 1.76) mg/d. N = 92.

Acupuncture for schizophrenia (Review)



#### acupoint cat - Sun 2005 (Continued)

	Combination therapy: could add artane, propranolol and benzodiazepine drugs and could not received any other antipsychotics.
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .
	Mental state <sup>2</sup> : PANSS; SAPS.
	Behaviour: leaving the study early.
	Adverse effects: TESS; skin infection.
	Unable to use:
	Adverse effects: lab test (kidney function; blood sodium; blood chloride; blood glucose; stool routine test) (reported no impact in both groups but no data).
	Lab test: EEG (not clinical outcome).
Notes	1. Assessment criteria (according to traditional criteria): recovery; marked improvement; improvement; no effect.
	2. The ratings were assessed before treatment and, 1 week after treatment, 2 weeks after treatment, 4 weeks after treatment and 6 weeks after treatment.

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Report stated - "using random sampling".
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Not used sham acupoint catgut treatment.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Unclear risk	This study was supported by the Health Department of Shanxi Province.

acupoint cat - Wang 1997		
Methods	Allocation: randomised. Blindness: not reported. Duration: 3 treatment courses.	

Acupuncture for schizophrenia (Review)

acupoint cat - Wang 1997 (Co	ntinued)	
Participants	Diagnosis: schizophrenia with auditory hallucinations (CCMD-2-R). N = 216.	
	Age: between 15 and 60 years (mean age (36.4 ± 2.1) years) (acupoint catgut treatment + antipsychotics group); between 20 and 58 years (mean age (35.7 ± 1.2) years) (antipsychotics group). Sex: 109 women and 107 men.	
	History: duration of illn ment + antipsychotics ; group). Setting: hospitalised pa	less between 1 and 8 years (average duration 2.3 years) (acupoint catgut treat- group); between 0.6 and 9.0 years (average duration 2.1 years) (antipsychotics atients.
	Country: China.	
	Inclusion criteria: had a 8-weeks treatment wit	a CCMD-2-R diagnosis of schizophrenia; auditory hallucinations still existed after h equivalent chlorpromazine dose 400 to 600 mg/d.
Interventions	<ol> <li>Acupoint catgut treatment + antipsychotics: acupoint catgut treatment (Tinggong [double]; opened mouth; 10 days as a treatment course; total 3 treatment courses) + antipsychotics (no further details).</li> <li>N = 108.</li> </ol>	
	2. Antipsychotics: no further details. N = 108.	
Outcomes	Mental state: no clinically important change in specific symptoms (auditory hallucinations) <sup>1</sup> .	
	Behaviour: leaving the study early.	
	Unable to use:	
	Adverse effects: local pain when eating (existed in acupoint catgut treatment + antipsychotics group but no data).	
Notes	1. Assessment criteria: marked improvement (auditory hallucinations disappeared within 10 days); im- provement (auditory hallucinations disappeared within 20 days); no effect (auditory hallucinations still existed after 30 days).	
	2. The equivalent chlorpromazine dose of two compared groups was no difference.	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into treatment group and control group ac- cording to admission order". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.

Blinding of participants and personnel (perfor- mance bias) All outcomes	Unclear risk	Not used; sham acupoint catgut treatment.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.

Acupuncture for schizophrenia (Review)



### acupoint cat - Wang 1997 (Continued)

Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.

Other bias Low risk Not obvious.

acupoint inj - Pan 200	2
Methods	Allocation: randomised. Blindness: assessor blind. Duration: 1 week washout period and 3 treatment courses. Follow-up: 2 years.
Participants	Diagnosis: schizophrenia (CCMD-2-R). N = 170. Age <sup>1</sup> : between 16 and 67 years (mean age (32.24 ± 10.40) years) (acupoint injection + chlorpromazine group); between 17 and 60 years (mean age (32.07 ± 10.17) years) (chlorpromazine group). History <sup>1</sup> : duration of illness between 3 and 240 months (average duration (54.72 ± 58.75) months) (acu- point injection + chlorpromazine group); between 3 and 304 months (average duration (65.81 ± 67.98)
	months) (chlorpromazine group). Sex: women and men. Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; BPRS > 36.
	Exclusion criteria: severe somatic and haemorrhagic diseases; acupuncture syncope and had aggres- sive behaviour.
	Follow-up: participants with effective treatment.
	N = 144/170. Age: between 16 and 60 years (mean age (31.59 ± 10.28) years(acupoint injection + chlorpromazine group); between 17 and 59 years (mean age (30.79 ± 9.61) years) (chlorpromazine group). Sex: 43 women and 101 men.
	History: duration of illness between 3 and 232 months (average duration (55.87 ± 59.76) months (acu- point injection + chlorpromazine group); between 3 and 301 months (average duration (53.02 ± 58.32) months) (chlorpromazine group).
Interventions	1. Acupoint injection + low dose chlorpromazine: acupoint injection <sup>*</sup> (Salviae Miltiorrhizae; acupoints choice according to the type of TCM <sup>**</sup> ) + low dose chlorpromazine (Wintermin; less than 300 mg/d; average dose (253.92 ± 42.25) mg/d <sup>1</sup> ). N = 105.
	2. Chlorpromazine: Wintermin; over 400 mg/d; average dose was (465.29 ± 72.92) mg/d <sup>1</sup> . N = 65.
	Combination therapy: dose could be adjusted within the provision range or added anticholinergic drugs when adverse reactions took place; could add benzodiazepine drugs when needed.
	Follow-up:
	1. Acupoint injection + chlorpromazine: Wintermin; maintained the same antipsychotics; average dose (149 $\pm$ 55) mg/d. N = 92/105.
	2. Chlorpromazine: Wintermin; maintained the same antipsychotics; average dose (160 $\pm$ 58) mg/d. N = 52/65.

Acupuncture for schizophrenia (Review)



acupoint inj - Pan 2002 (Contir	nued)	the same as treatment period
	* Acupoint injection: tw val between two course	vo side acupoints in turn; once a day; 10 days as a treatment course; 7-day inter- es.
	** Acupoints choice aco	cording to type of TCM:
	Type of phlegm-fire att	acking upwards: Shangqiu, Fenglong, Yangjiao, Ququan, Baihui.
	Type of internal retenti Baihui.	on of phlegm and dampness; Sanjingjiao, Shangqiu, Fenglong, Yanglingquan,
	Type of Qi stagnation a	nd Blood stasis: Ligou, Benshen, Yangjiao, Ququan, Baihui.
	Type of Yin deficiency a	and fire excess: Shaohai, Zhizheng, Zhubing, Feiyang, Baihui.
	Type of Yang deficiency	r: Sanyingjiao, Zusanli, Dazhong, Feiyang, Baihui.
Outcomes	Mental state: BPRS <sup>2,3</sup> .	
	Behaviour: leaving the	study early.
	Adverse effects: TESS <sup>4</sup> .	
	Follow-up:	
	Global state: relapse.	
	Mental state:BPRS <sup>5</sup> (BF	PRS score was assessed continuously during follow-up).
Notes	1. Data reported from o	only 160 participants.
	2. The rating was asses	sed before treatment and after each treatment course.
	3. Another assessment 50%); improvement (≥	standard (according to reduced rate): recovery (≥ 75%); marked improvement (≥ 25%); no effect (< 25%).
	4. The rating was asses	sed after each treatment course.
	5. The rating was asses	sed before follow-up, one year and two years; we used data of two years.
	6. Contact made with a had retired and author those outcomes contai	uthor as there was little difference between two references. Author Yufeng Pan Yuying Pan clarified that the two references referred to the same study. For ning different data from the two papers we did not extract the data.
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera-	Unclear risk	Report stated - "randomly divided into the two groups". No further details.

Not stated.

Sham acupoint injection not used.

Report stated - "the assessors were blind to the drug allocation".

Acupuncture for schizophrenia (Review)

tion (selection bias)

(selection bias)

mance bias) All outcomes

Allocation concealment

**Blinding of participants** 

and personnel (perfor-

Blinding of outcome as-

sessment (detection bias)

Copyright @ 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Unclear risk

High risk

Low risk



### acupoint inj - Pan 2002 (Continued) All outcomes

Incomplete outcome data (attrition bias) Outcomes	High risk	Ten participants left the study early during the study and the follow-up period.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	High risk	There were inconsistencies in the data between the 2 reports of the same study.

### acupoint inj - Wang 2000

Methods	Allocation: randomised. Blindness: assessor blind. Duration: 2 weeks.	
Participants	Diagnosis: schizophrenia with auditory hallucinations (CCMD-2-R). N = 90.	
	Age: between 18 and 59 group); between 18 and Sex: 31 women and 59	9 years (mean age (39.54 ± 10.93) years) (acupoint injection + antipsychotics d 58 years (mean age (38.27 ± 9.78) years) (antipsychotics group). men.
	History: duration of illness (12.70 ± 8.31) years (acupoint injection + antipsychotics group); (11.5 ± 8.31) years (antipsychotics group). Setting: hospitalised patients.	
	Country: China.	
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; existed all kinds of auditory hallucina- tions and the scores of the twelfth item of BPRS ≥ 3; auditory hallucinations did not disappear after 2 months systemic treatment with one antipsychotics.	
	Exclusion criteria: organic diseases and other hallucinations.	
Interventions	1. Acupoint injection + antipsychotics: acupoint injection (clonazepam 1.0 mg; Tinggong [double], opened mouth, once two days, 7 times) + antipsychotics (remained previous antipsychotics treatment; equivalent chlorpromazine dose (685 ± 240) mg/d). N = 45.	
	2. Antipsychotics: remained previous antipsychotics treatment; equivalent chlorpromazine dose (633 ± 247) mg/d. N = 45.	
Outcomes	Mental state: BPRS (the scores of the twelfth item).	
	Behaviour: leaving the study early.	
Notes		
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly divided into experimental group and control group using draw lots method".

Acupuncture for schizophrenia (Review)

### acupoint inj - Wang 2000 (Continued)

Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham acupoint injection not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - " two doctors with extensive clinical experience and not at- tending the trial assessed outcomes before treatment and 2 weeks after treat- ment".
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	High risk	One author worked for drug industry.

### acupoint inj - Yang 2000

Methods	Allocation: randomised. Blindness: not reported. Duration: half a year.
Participants	Diagnosis: schizophrenia with auditory hallucinations (CCMD-2). N = 64.
	Age: mean age 30.7 years (acupoint injection + antipsychotics group); 28.8 years (antipsychotics group). Sex: 10 women and 54 men.
	History: average duration of illness 5.2 years (acupoint injection + antipsychotics group) and 5.8 years (antipsychotics group).
	Country: China.
	Inclusion criteria: had CCMD-2-R diagnosis of schizophrenia; > 2 hospitalisations; systemic treatments of multiple antipsychotics but persistent auditory hallucinations; after other treatments (acupuncture and Chinese herbs) persistent auditory hallucinations or reappeared; current auditory hallucinations > 8 weeks; BPRS < 30.
Interventions	1. Acupoint injection + antipsychotics: acupoint injection (sulpiride 50 mg; Tinggong [double], once a day, 5 times as a treatment course, 2 intermittent treatment courses each month, total half a year) + antipsychotics (remained antipsychotics treatment; equivalent chlorpromazine dose (668.4 ± 221.6) mg/d). N = 34.
	2. Antipsychotics: remained antipsychotics treatment (except some cases reduced dose or added ar- tane for extrapyramidal reaction); equivalent chlorpromazine dose (651 ± 20.84) mg/d. N = 30.
Outcomes	Mental state: BPRS <sup>1</sup> ; no clinically important change in specific symptoms (auditory hallucinations) <sup>2</sup> .
	Behaviour: leaving the study early.
Notes	1. The rating was assessed 1 week before treatment and, 3 months and half a year after treatment.

Acupuncture for schizophrenia (Review)



#### acupoint inj - Yang 2000 (Continued)

2. Criteria: recovery (auditory hallucinations disappeared completely; did not reappear about half a year); improvement (times of auditory hallucinations reduced; had insight of auditory hallucinations; did not effect daily life; or auditory hallucinations appeared occasionally after disappeared); no effect (auditory hallucination still existed or appeared frequently after disappeared).

#### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham acupoint injection not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

#### EACT - Xue 1987

Methods	Allocation: randomised. Blindness: not reported. Duration: a treatment course.
Participants	Diagnosis: schizophrenia. N = 68.
	Age: between 19 and 37 years (electric acupuncture convulsive therapy group) and between 17 and 48 years (electroconvulsive therapy group); most of them were middle-aged patients. Sex: 23 women and 45 men.
	Country: China.
Interventions	1. Electric acupuncture convulsive therapy: Renzhong and Baihui; average electricity consumption 1.27 Joule/331 times; once every two days; 12 times as a treatment course. N = 34.
	2. Electric convulsive therapy: electrode position of two temporal; average electricity consumption 34.97 Joule/286 times; once every two days; 12 times as a treatment course. N = 34.
Outcomes	Behaviour: leaving the study early.
	Adverse effects: back pain; spinal fracture <sup>1</sup> .

Acupuncture for schizophrenia (Review)

#### EACT - Xue 1987 (Continued)

	Unable to use:		
	Physical exam: tendon reflexes; patellar clonus (not clinical outcomes).		
Notes	1. Twelve patients suffered a spinal fracture during the treatment period and their data could not be used.		
	2. Authors compared three groups (electric acupuncture convulsive therapy group, electroconvulsive therapy group and grand mal epilepsy group) but only patients with schizophrenia randomly divided into electric acupuncture convulsive therapy group and electroconvulsive therapy group.		

### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Author compared two groups and used electrodes with different positions.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### electro - Chen 2006

Methods	Allocation: randomised. Blindness: not reported. Duration: 6 weeks.
Participants	Diagnosis: post-schizophrenic depression <sup>1</sup> (CCMD-3).
	N = 67. Age: mean age (30.2 ± 11.0) years (electroacupuncture + antipsychotics group) and (2.9 ± 10.7) years <sup>2</sup> (antipsychotics group). Sex: 27 women and 40 men.
	History: average duration of illness (5.6 $\pm$ 2.9) years (electroacupuncture + antipsychotics group) and (4.9 $\pm$ 3.8) years (antipsychotics group).
	Setting: both inpatients and outpatients.

Acupuncture for schizophrenia (Review)

### electro - Chen 2006 (Continued)

	Country: China.		
Interventions	1. Electroacupuncture + antipsychotics: electroacupuncture <sup>*</sup> (Baihui and Yingtang) + antipsychotics (remained previous medication). N = 33.		
	2. Antipsychotics: remained previous medication. N = 34.		
	* Electroacupuncture: continuous wave; 2-4 Hz, 50 minutes, once a day.		
Outcomes	Mental state: HAMD <sup>3,4</sup> .		
	Behaviour: leaving the study early.		
	Adverse effects: TESS <sup>3</sup> .		
Notes	1. Diagnosis of post-schizophrenia depression: diagnosed with schizophrenia in the last 1 year; symp- toms of depression appeared when the condition improved but not cured; the depression continued at least two weeks; HAMD > 20.		
	2. Although the author reported that there was no significant age difference between the two groups the table appears to contain a typographical error in the antipsychotics' group.		
	3. The rating was assessed before treatment and at 2 weeks, 4 weeks and 6 weeks after treatment.		
	4. Another assessment standard (according to reduced rate): recovery (≥ 75%); marked improvement (≥ 50%); improvement (≥ 50%); no effect (< 25%, reduced rate = [total scores before treatment-total scores after treatment]/total scores before treatment*100%).		

### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.



electro - Chen 2008	
Methods	Allocation: randomised. Blindness: not report. Duration: 12 weeks.
Participants	Diagnosis: schizophrenia with negative symptoms (type II schizophrenia) (CCMD-3; Andreasen's diag- nosis standard of type II schizophrenia) .
	N = 62. Age: between 19 and 54 years (mean age (27 ± 13) years) <sup>1</sup> (electroacupuncture + aripiprazole group); between 17 and 60 years (mean age (28 ± 15) years) (aripiprazole group). Sex: 22 women and 38 men <sup>1</sup> .
	History: duration of illness between 3 and 24 years <sup>1</sup> (electroacupuncture + aripiprazole group); be- tween 2 and 19 years (aripiprazole group).
	Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-3 and a type II diagnosis of schizophrenia; with mainly negative symp- toms and PANSS > 60, the scores of negative factors > 30; age between 16 and 60 years and duration of illness more than 1 year; liver and kidney function test, lipids test, blood glucose test, ECG, blood rou- tine test, urine routine test, and stool routine test normal; patients or family members gave their con- sent.
	Exclusion criteria: combined with other diseases; had serious physical illnesses or brain organic dis- eases or epilepsy and alcohol or drug abuse; pregnant or lactating women; with similar drug allergic history and unable to adapt electroacupuncture history.
Interventions	1. Electroacupuncture + aripiprazole: electroacupuncture <sup>*</sup> (Baihui and Neiguan [double] group or Shuigou and Sanyingjiao [double] group) + aripiprazole (Bosiqing; initial dosage 5 mg/d; gradually added to the therapeutic dosage 10 to 30 mg/d within two weeks; average dosage (18.4 ± 6.2) mg/d). N = 32.
	2. Aripiprazole: Bosiqing; initial dosage 5 mg/d; gradually added to the therapeutic dosage 10 to 30 mg/ d within two weeks; average dosage (20.1 ± 4.3) mg/d. N = 30.
	Washout period: those treated with other antipsychotics or mood stablisers needed 2-weeks' washout period.
	Combination therapy: could not receive any other antipsychotics during the observation period; symp- tomatic treatment drugs could be used according to patient's condition when with adverse drug reac- tions and could receive benzodiazepines when with poor sleep.
	* Electroacupuncture: used two acupoints groups in turn; once for 45 minutes; once a day; five days a week (except Saturday and Sunday); 12 weeks as a treatment course.
Outcomes	Mental state: PANSS <sup>2,3</sup> .
	Behaviour: leaving the study early.
	Adverse effects: TESS <sup>2</sup> (mainly reported insomnia; myotonia; tremor; akathisia; blurred vision; sweat- ing; headache; tachycardia).
	Unable to use:
	Adverse effects: lab test (kidney function test, lipids test, blood glucose test, stool routine test) (report- ed no obvious abnormal before or after treatment in both groups and no data).
Notes	1. Two patients in the electroacupuncture + aripiprazole group left the study early and author reported the age, sex and duration of illness of this group excluding those two patients.

Acupuncture for schizophrenia (Review)

#### electro - Chen 2008 (Continued)

Librarv

2. The ratings were assessed before treatment and at 2 weeks, 4 weeks, 8 weeks and 12 weeks after treatment.

3. Another assessment standard (according to reduced rate): recovery (≥ 75%); marked improvement (51% to 75%); improvement (25% to 50%); no effect (< 25%, reduced rate = [scores before treatment-scores after treatment]/scores before treatment\*100%).

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Report stated - "random systematic sampling according to admission order".
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	High risk	Two participants (electroacupuncture + aripiprazole group) left the study ear- ly.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

#### electro - Cheng 2009

Methods	Allocation: randomised. Blindness: patient and assessor blind.	
	Duration: 4-week baseline evaluation and risperidone run-in phase + 6 weeks treatment period.	
Participants	Diagnosis: schizophrenia with auditory hallucinations (DSM IV) .	
	N = 60. Age: mean age (31.20 ± 5.79) years (electroacupuncture + risperidone group) and (28.97 ± 5.49) years (sham electroacupuncture + risperidone group). Sex: 32 women and 28 men.	
	History: average duration of auditory hallucinations (10.13 $\pm$ 2.89) years (electroacupuncture + risperi- done group) and (9.03 $\pm$ 2.94) years (sham electroacupuncture + risperidone group).	
	Setting: hospitalised patients.	
	Country: China.	
	Inclusion criteria: were 18-60 years of age; had a DSM IV diagnosis of schizophrenia using Structured Clinical Interview for DSM IV; had previously been treated with risperidone following documented	

Acupuncture for schizophrenia (Review)

electro - Cheng 2009 (Continued	) 	
	dose; demonstrated a c of risperidone (three or tressing auditory halluc informed consent.	wo antipsycholics were administered for an adequate duration with sufficient locumented failure to show a satisfactory clinical response to an adequate trial more months of at least 4 mg/day of oral risperidone); had persistent and dis- inations, evidenced by a score of 11 or more on the total of PSYRATS-AH; gave
	Exclusion criteria: ment vere and unstable phys acupuncture (to maxim	al retardation; seizure disorder; substance abuse/dependence; pregnant; se- ical illnesses; danger of attack or extreme agitation; or previously received ise blinding).
Interventions	1. Electroacupuncture - and Sanyijiao) + risperio on a stable dose of risp throughout the study; t	Frisperidone: electroacupuncture <sup>*</sup> (Tinggong, Tinghui, Yifeng, Daling, Neiguan done (before initiation of the baseline observation period all patients had been eridone for at least 4 weeks and baseline doses of risperidone remained stable he average initial risperidone dosage (5.15 ± 0.46) mg/d). N = 30.
	2. Sham electroacupun sponding acupoint) + ri been on a stable dose o stable throughout the s	cture + risperidone: sham electroacupuncture (20 mm away from each corre- speridone (before initiation of the baseline observation period all patients had if risperidone for at least 4 weeks and baseline doses of risperidone remained tudy; the average initial risperidone dosage (5.22 ± 0.47) mg/d). N = 30.
	Combination therapy: l mood stabilisers and ar	orazepam (< 4 mg) was allowed to counteract sleep problems; antidepressants, ntipsychotic drugs other than risperidone were not allowed during the study.
	* Electroacupuncture: r the handles of the need cal input via an electroa mA); for 20 minutes; 5 t	needled bilaterally; depth 15to 30mm; once the 'De-Qi' sensation was elicited, lles inserted into Tinggong, Tinghui and Yifeng were connected to an electri- acupuncture machine (sparse dense wave; frequency 2 to 10 Hz; intensity 2 to 3 imes a week; total 30 times (6 weeks).
	** Sham electroacupun near Tinggong, Tinghui cal current; for 20 minu ture procedure and the elicited).	cture: depth less than 5 mm; the handles of the needles inserted into points and Yifeng were connected to the electroacupuncture machine with no electri- tes; 5 times a week; total 30 times (6 weeks) (simulated a real electroacupunc- acupuncturist did not manipulate the needles and no 'De-Qi' sensation was
Outcomes	Mental state: PSYRATS-AH <sup>1,2</sup> ; PANSS <sup>1</sup> .	
	Behaviour: leaving the study early.	
Notes	1.The ratings were assessed at baseline, 2 weeks, 4 weeks and 6 weeks.	
	2. Another assessment standard: (according to reduction scores from baseline): a treatment response (≥ 20%).	
	3. This study is a part of information from autho	Bai's study (see the reference) and they only reported this study (obtained the or Jing Chen).
	4. Author analysed data	using intention-to-treat method.
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly assigned to either the real electroacupuncture group or the sham electroacupuncture group by the SAS program".
Allocation concealment (selection bias)	Low risk	Report stated - "maintained using opaque sealed envelopes".

Low risk Report stated - "blinded to the treatment allocation"; the acupuncturist was Blinding of participants "instructed not to communicate with the patients and the clinical investigators"; the participants were "acupuncture-naive patients" and "there was al-

Acupuncture for schizophrenia (Review)

and personnel (perfor-

mance bias)



### electro - Cheng 2009 (Continued)

All outcomes		so limited contact between the study participants and restricted conversation between acupuncturist and participants during treatment".
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - "blinded to the treatment allocation" and the acupuncturist was "instructed not to communicate with the patients and the clinical investigators".
Incomplete outcome data (attrition bias) Outcomes	Low risk	Though seven participants left the study early the author analysed data using intention-to-treat method.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Unclear risk	This study was supported by the Stanley Medical Research Institue (SMRI), Chevy Chase, Maryland.

electro - Cui 2000	
Methods	Allocation: randomised. Blindness: not reported.
	Duration: 6 weeks.
Participants	Diagnosis: schizophrenia (CCMD-2-R) .
	N = 60. Age: between 20 and 50 years (mean age (35.1 ± 7) years). Sex: 48 women and 12 men.
	History: duration of illness between 5 and 10 years.
	Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; without severe heart, liver and kidney dysfunction or other severe somatic diseases; without chlorpromazine allergic history.
Interventions	1. Electroacupuncture + chlorpromazine: electroacupuncture <sup>*</sup> (acupoint choice according to different symptoms) + chlorpromazine (100 to 300 mg/d). N = 30.
	2. Chlorpromazine: 400 to 500 mg/d. N = 30.
	Combination therapy: not combined with any other antipsychotics; could received symptomatic treat- ment when with extrapyramidal reactions, tachycardia or elevated transaminase.
	* Electroacupuncture: with mainly positive symptoms used Xinshu, Ganshu, Pishu, Shenmen, Feng- long; with mainly negative symptoms used Dazhui, Fengfu, Neiguan, Fenglong; with both positive and negative symptoms added or removed acupoints according to the symptoms; refusing to eat added Hegu and Zusanli; not speaking added Lianquan and Yamen; excitement and hyperactivity added Ren- zhong and Quchi; intensity according to patients feeling; frequency 20 to 40 times/second; for 30 min- utes; once two days; 20 times as a treatment course.
Outcomes	Mental state: BPRS <sup>1,2</sup> .
	Behaviour: leaving the study early.

Acupuncture for schizophrenia (Review)

electro - Cui 2000 (Continued)	Adverse effects: TESS <sup>3</sup> (mainly reported extrapyramidal symptoms; tachycardia; dry mouth; blurred vi- sion; sleepiness).
	Others <sup>4</sup> : adding medication.
Notes	1.The rating was assessed before treatment and, 2 weeks, 4 weeks and 6 weeks after treatment.
	2. Another assessment standard (according to reduced rate): improvement (≥ 30%); marked improve- ment (≥ 50%); recovery (≥ 80%); no effect (< 30%).
	3. The ratings were assessed at 2 weeks, 4 weeks and 6 weeks after treatment.
	4. Author reported this outcome.

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Report state - "randomly divided into two groups according to age and gender matching". Contacted the author who said she used coin-tossing method .
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

electro - Ding 2005	
Methods	Allocation: randomised. Blindness: not reported. Duration: 2 months.
Participants	Diagnosis: chronic schizophrenia (CCMD-3).
	N = 50. Age: between 47 and 60 years (mean age (57.4 ± 5.71) years). Sex: 50 men and no women.
	History: duration of illness between 19 and 39 years (average duration (32.25 ± 6.31) years); duration of continued hospitalisation between 11 and 22 years (average duration (18.17 ± 7.35) years); received an-tipsychotics more than 10 years.

Acupuncture for schizophrenia (Review)

electro - Ding 2005 (Continued)	Setting: hospitalised p	atients.
	Country: China.	
Interventions	1. Electroacupuncture long [double], Taichon N = 25.	+ antipsychotics: electroacupuncture <sup>*</sup> (Tanzhong, Zhongwan, Shenmen, Feng- g [double], Neiguan [double]) + antipsychotics (remained previous medication).
	2. Antipsychotics: rema	ained previous medication. N = 25.
	* Electroacupuncture: excess syndrome; conr ed; pulse current; 5 min	reinforcing-reducing method for deficiency syndrome and reducing method for nected needles to the electroacupuncture machine once the sensation was elicit- nutes; once two days.
Outcomes	Mental state: BPRS; PA	NSS.
	Behaviour: leaving the	study early.
Notes	1. Contact made with t	he author, Fenggang Li, and no participants left the study early.
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into treatment group and control group". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Not used sham electroacupuncture and doctors often told patients (elec- troacupuncture + antipsychotics group) the importance and treatment mecha- nism of acupuncture.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Unclear risk	Participants were men and unsure if gender selection bias occurred and the study lasted longer than 20 years.

### electro - Wang 2005

Methods	Allocation: randomised. Blindness: not reported. Duration: 8 weeks.
Participants	Diagnosis: schizophrenia with negative symptoms (type II schizophrenia) (CCMD-3; Andreasen's diag- nosis standard of type II schizophrenia).

Acupuncture for schizophrenia (Review)

electro - Wang 2005 (Continued)	N = 75.		
	Age: mean age (41.7 ± 1 done group). Sex: 41 women and 34 r	0.6) years (electroacupuncture + risperidone group); (39.3 ± 9.2) years (risperi- men.	
	History: duration of illn years (risperidone grou risperidone group) and Setting: hospitalised pa	ess (11.7 $\pm$ 6.5) years (electroacupuncture + risperidone group) and (13.2 $\pm$ 7.4) p); average number of hospitalisations (4.2 $\pm$ 2.5) times (electroacupuncture + (3.9 $\pm$ 2.2) times (risperidone group). attents.	
	Country: China.		
	Inclusion criteria: had a schizophrenia; PANSS > years; patients receivin washout period and tot	CCMD-3 diagnosis of schizophrenia; had an Andreasen's diagnosis of type II 60; the scores of negative factors > 30; women and men; age between 18 and 55 g any other antipsychotics, mood stabilisers or antidepressants needed 2 weeks cal scores of PANSS decreased < 20.	
	Exclusion criteria: had o chotic diseases; with sin lactating women.	cerebral organic diseases or severe physical illnesses; combined with other psy- milar drugs allergic history; drug or alcohol dependence or abuse; pregnant or	
Interventions	1. Electroacupuncture - [double], Neiguan [dou dose 3 to 6 mg/d in 10 c ± 1.4) mg/d). N = 40.	+ risperidone: electroacupuncture <sup>*</sup> (Baihui, Shangxing, Yingtang, Sanyingjiao ble]) + risperidone (Weisitong; initial dose 1 mg/d and added to therapeutic lays according to patient's conditions and adverse reactions; average dose (4.9	
	2. Risperidone: Weisitor cording to patient's cor	ng; initial dose 1 mg/d and added to therapeutic dose 3 to 6 mg/d in 10 days ac- nditions and adverse reactions; average dose ( $5.3 \pm 1.2$ ) mg/d. N = 35.	
	Combination therapy: o add benzodiazepine dr	could add artane, propranolol when extrapyramidal reactions appeared; could ugs to treat patients with pool sleep.	
	* Electroacupuncture: f week; 20 times as a trea comfortable symptoms and Qi.	requency 20 to 40 times/minute; intermittent wave; for 45 minutes; 5 times a atment course; added Zhongwan and Zusanli when with gastrointestinal un- and added Fenglong when with symptoms of the type of stagnation of phlegm	
Outcomes	Mental state: PANSS <sup>1,2</sup> .		
	Behaviour: leaving the	study early.	
	Adverse effects: TESS <sup>1</sup> ness; constipation; weig liver function and kidne	(main: extrapyramidal symptoms; dry mouth; insomnia; blurred vision; dizzi- ght gain; nausea and vomiting); lab test (blood routine test; urine routine test; ey function; and ECG).	
	Others <sup>3</sup> : adding medica	ation.	
Notes	1. The ratings were asse	essed at the end of 0 week, 4 weeks and 8 weeks.	
	2. Another assessment (51% to 75%); improver	standard (according to reduced rate): recovery (≥ 75%); marked improvement ment (25% to 50%); no effect (< 25%).	
	3. Author reported this	outcome.	
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into experimental group and control group". No further details.	

Acupuncture for schizophrenia (Review)

### electro - Wang 2005 (Continued)

Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### electro - Xiong 2010

Methods	Allocation: randomised. Blindness: assessor blind. Duration: 8 weeks.
Participants	Diagnosis: refractory schizophrenia (CCMD-3, the defined standard of refractory schizophrenia <sup>1</sup> ). N = 80.
	Age: mean age (29.4 ± 11.3) years (electroacupuncture + low dose clozapine group); (28.1 ± 12.2) years (clozapine group). Sex: 34 women and 46 men.
	History: average duration of illness (35.2 ± 12.2) months (electroacupuncture + low dose clozapine group); (36.3 ± 13.2) months (clozapine group). Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-3 diagnosis of schizophrenia; had a diagnosis of refractory schizophre- nia; age between 14 and 60 years; women and men; patients already receiving antipsychotics needed 7 days' washout period; gave informed consent.
	Exclusion criteria: had severe heart, liver and kidney primary diseases.
Interventions	1. Electroacupunture + low dose clozapine: electroacupuncture <sup>*</sup> (Baihui and Taiyang [double]) + low dose clozapine (initial dose 50 mg/d; added to 100 to 150 mg/d within 1 week). N = 40.
	2.Clozapine: initial dose 50 to 100 mg/d, added to 200 to 500 mg/d within 1 week. N = 40.
	* Electroacupuncture: 10 voltage; continuous wave; frequency 60Hz; current intensity was limited when neck and facial muscles cramped and breath-hold and hypoxia appeared; firstly continued to stimulate strongly 3 to 4 seconds then reduced the strength and frequency quickly, after patient's breath and face returned to normal and continued 30 to 60 seconds stimulated again; continued to stimulate 8 to 10 times; 3 times a week, total 8 weeks.
Outcomes	Mental state: PANSS <sup>2,3</sup> .

Acupuncture for schizophrenia (Review)

### electro - Xiong 2010 (Continued)

Behaviour: leaving the study early.

	Adverse effects: TESS <sup>4</sup> .
Notes	<ol> <li>The defined standard of refractory schizophrenia: no effect after receiving at least 2 or more different antipsychotic chemical drugs with enough doses and enough treatment courses; no effect after receiv- ing electric shock 7 to 12 times enough treatment courses' treatments; PANSS ≥ 60.</li> </ol>
	2. The rating was assessed before treatment and, 2 weeks, 4 weeks and 8 weeks after treatment.
	3. Another assessment standard (according to reduced rate): recovery (≥80%); marked improvement (≥ 50%, < 80%); improvement (≥ 30%, < 50%); no effect (< 30%, reduced rate = [scores before treat- ment-scores after treatment]/scores before treatment*100%).
	4. The rating was assessed 8 weeks after treatment.

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomised into" two groups. No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - "two doctors who were blind the drug allocation assessed out- comes".
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

electro - Yao 2006	
Methods	Allocation: randomised. Blindness: not reported.
	Duration: 8 weeks.
	Follow-up: 6 months (effective case).
Participants	Diagnosis: schizophrenia (CCMD-3) .
	N = 90. Age: between 18 and 60 years (mean age (31 ± 12) years) (electroacupuncture + clozapine group); be- tween 18 and 60 years (mean age (29 ± 13) years) (clozapine group).

Acupuncture for schizophrenia (Review)



electro - Yao 2006 (Continued)	Sex: 40 women and 50 r	nen.
	History: duration of this troacupuncture + cloza (clozapine group).	s episodes between 1 and 16 months (average duration (8 $\pm$ 5) years) (elecpine group), between 1.5 and 15 months (average duration (9 $\pm$ 3) months)
	Type: paranoid type 310 positive family history ( ed type 12 cases, hebep	cases, undifferentiated type 9 cases, hebephrenic type 5 cases and 8 cases with electroacupuncture + clozapine group); paranoid type 29 cases, undifferentiat- phrenic 4 cases and 10 cases with positive family history (clozapine group).
	Setting: hospitalised pa	tients.
	Country: China.	
	Inclusion criteria: had a last two weeks.	CCMD-3 diagnosis of schizophrenia; not received any antipsychotics within the
	Exclusion criteria: seven pendence; pregnant or	re heart, brain, blood vessels and other physical illnesses; drug or alcohol de- lactating women.
Interventions	1. Electroacupuncture + clozapine: electroacupuncture <sup>*</sup> (Baihui, Fenglong, Houxi, Ganshu) + clozapine (total dose 200-300 mg/d; twice a day). N = 45.	
	2. Clozapine: total dose	200 to 300 mg/d; twice a day. N = 45.
	* Electroacupuncture: s course; total 2 treatmer	parse dense wave; for 30 minutes; once a day; 20 to 30 times as a treatment nt courses.
Outcomes	Mental state: PANSS <sup>1.2</sup> .	
	Behaviour: leaving the	study early.
	Adverse effects: TESS <sup>1,3</sup>	3.
	Follow-up:	
	Mental state: PANSS <sup>2,4</sup> .	
Notes	1. The ratings were asse ment.	essed before treatment and, 2 weeks, 4 weeks, 6 weeks and 8 weeks after treat-
	2. Another assessment (60% to 94%); improver	standard (according to reduced rate): recovery (> 95%); marked improvement nent (30-59%); no effect (< 30%).
	3. Lab test included in T	ESS was tested before treatment and after treatment.
	4. Effective cases included recovery cases, marked improvement cases and improvement cases. Effec- tive cases were assessed 6 weeks of follow-up period (converted to dichotomous data).	
Risk of bias		
Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Low risk	Reported state - " randomly divided into two group using draw lots method".
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias)	High risk	Sham electroacupuncture not used.

Acupuncture for schizophrenia (Review)



Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	High risk	No participants left the study early but only followed up effective cases and six participants left the study early at follow-up period.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

electro - Zhang 1987	
Methods	Allocation: randomised. Blindness: not reported. Duration: 20 days.
Participants	Diagnosis: schizophrenia.
	First stage: N = 182. Age: between 17 and 43 years (electroacupuncture + Dang Gui Cheng Qi Tang group); between 17 and 51 years (decoction of herbs group); between 17 and 48 years (electroacupuncture group); between 16 and 48 years (chlorpromazine group). Sex: 97 women and 85 men.
	History: mostly first admissions. Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: as for readmitted patients only those who experienced marked improvement or cure at the end of the previous course of treatment were included. Exclusion criteria: those whose illness had lasted over two years; weak or pregnant; had symptoms of deterioration; physically frail or had a history of peptic ulcer, cardiac or renal disease; those who showed rapid remission of symptoms on admission.
	Second stage:
	N = 99. Age: under 20 years 16 cases; between 21 and 30 years 52 cases; between 31 and 40 years 18 cases; be- tween 41 and 50 years 10 cases; over 50 years 3 cases. Sex: all men and no women.
	History: duration of illness within 2 years 66 cases and over 2 years 33 cases; first admissions 32 cases, second admissions 21 cases and more than 3 hospitalisation times 9 cases. Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: as for readmitted patients only those who experienced marked improvement or cure at the end of the previous course of treatment were included.
	Ecliusion criteria: those whose illness had lasted over two years were included; others were the same as the first stage.

Acupuncture for schizophrenia (Review)

Copyright @ 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

electro - Zhang 1987 (Continued)			
Interventions	First stage:		
	1. Electroacupuncture + Dang Gui Cheng Qi Tang: electroacupuncture <sup>*</sup> (Yifeng, Tinggong, Tounjie, Chengling, Linqi, Baihui, Dingshen) + Dang Gui Cheng Qi Tang <sup>**</sup> . N = 49.		
	2. Decoction of herbs: Dang Gui Cheng Qi Tang <sup>**</sup> . N = 45.		
	3. Electrocupuncture <sup>*</sup> : Yifeng, Tinggong, Tounjie, Chengling, Linqi, Baihui, Dingshen. N = 43.		
	4. Chlorpromazine (Thorazine): any dose when necessary; 300-600 mg/day; not combined with other antipsychotics. N = 45.		
	Second stage:		
	1. Electroacupuncture + Dang Gui Cheng Qi Tang: electroacupuncture <sup>*</sup> (Yifeng, Tinggong, Chengling and Tounie) + Dang Gui Cheng Qi Tang <sup>**</sup> . N = 26.		
	2. Electroacupuncture + Dang Gui Cheng Qi Tang (with additives): electroacupuncture <sup>*</sup> (Yifeng, Tinggong, Chengling and Tounie) + Dang Gui Cheng Qi Tang <sup>**</sup> (with additives <sup>***</sup> ). N = 25.		
	3. Electroacupuncture <sup>*</sup> : Tounie, Chengling and Linqi. N = 23.		
	4. Electroacupuncture <sup>*</sup> : Yifeng, Tinggong, Chengling and Tounie. N = 25.		
	* Electroacupuncture: successive waves of 120cycles (Hz)/sec and 500 μsec pulse width; the intensity of stimulation varied with the individual and could be generalised into three grades (strong stimulation, moderate stimulation and mild stimulation); twice a day; adjustment of treatment depended upon the condition of the patient.		
	** Dang Gui Cheng Qi Tang: Radix Angelicae Sinensis 30 g, Radix et Rhizoma Rhei 30 g, Natrii Sulfas 15 g, Poncirus Trifoliata (L) Raf 12 g and Fruetus Trichosanthis 150 mL; all herbs except the Natrii Sulfas were decocted into a 100 mL solution, then the Natrii Sulfas was added and dissolved in it; usually 50 mL two times a day but the dose maybe increased to 150 to 200 mL daily when necessary.		
	*** Additional herbs of Dang Gui Cheng Qi Tang (with additives): Swmen Persicae, Radix Curcumae, Radix Paeoniae Rubrae, Radix Bupleuri, Radix Scutellariae, Flos Carthami, Rhizoma Ligustici Chuanx- iong, Radix Srephaniae tetrandrae, Radix Ledebouriellae, Poria, Radix Polygalae, Fructus Ziziphi Ju- jubae, Radix Rehmanniae, Rhizoma Acori Graminei, Os Draconis, Concha Osreeae, Haematitum, Herba Leonuri, Parata, Cortex Cinnamoni, Rhizoma Coptidis; according to the different condition of patient.		
	Combination therapy: 10% chloral hydrate 10 to 20 mL orally, hyminal 0.1 to 0.2 g orally, phenobarbital 0.03 to 0.1 g or paraldehyde 4 to 5 mL orally or intramuscularly for patient who could not fall asleep at night and could not to be used for a long time.		
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .		
	Behaviour: leaving the study early.		
	Unable to use;		
	Adverse effects: reported electroacupuncture relative adverse effects (holding breath, facial cyanosis, arrhythmia, transient increase of blood pressure, injury of teeth, tongue and lips, epileptic attacks with strong stimulation) and TCM relative adverse effects (diarrhoea) existed but no data.		
Notes	1. Assessment criteria: recovery (disappearance of all symptoms); marked improvement (50% relief of symptoms with moderate insight); mild improvement (25% relief of symptoms with no insight); no effect (no change in symptoms).		
	2. Only included the first stage of the study and could not include the second stage of the study because of interventions.		
Risk of bias			

Acupuncture for schizophrenia (Review)

\_



### electro - Zhang 1987 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "divided into groups randomly". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture or any other dummy treatments not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

electro - Zhang 1993	
Methods	Allocation: randomised. Blindness: not reported. Duration: 8 weeks .
Participants	Diagnosis: refractory schizophrenia (CCMD-2-R). N: unclear <sup>1</sup> .
	Age: both references reported mean age.
	History: both references reported average duration of illness. Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; after at least 2 times hospitalisations and systemic treatments of multiple antipsychotics most of the psychiatric symptoms disappeared but au- ditory hallucinations appeared repeatedly; after other treatments (acupuncture and Chinese herbs) auditory hallucinations still existed or reappeared after disappeared; auditory hallucinations this time continued more than 8 weeks; BPRS < 30.
Interventions	1. Electroacupuncture + antipsychotics: electroacupuncture <sup>*</sup> (CCEA; group one [Yingtang and Baihui] and group two [Shenting and Yamen]) + antipsychotics <sup>**</sup> (remained unchanged doses and kinds of medicines).
	2. Antipsychotics**: remained unchanged doses and kinds of medicines.
	* Electroacupuncture: both groups were alternately used; sine-wave; frequency 12, 10, 8, 6 Hz respec- tively for 10, 10, 10 and 15 minutes; frequency of fundamental waves 250 Hz and 750 Hz; maxim voltage 2 to 9 V; for 45 minutes; once a day; 8 weeks as a treatment course.

Acupuncture for schizophrenia (Review)

#### electro - Zhang 1993 (Continued)

	** From the date of two months before dividing into two groups to the end of the study the doses and kinds of medicines taken remained unchanged.	
Outcomes	Mental state <sup>2</sup> : SANS; SAPS.	
	Unable to use:	
	Global state: no clinically important change in global state <sup>3</sup> (data could not to be used).	
	Mental state: BPRS (data could not to be used).	
	Behaviour: leaving the study early (data could not to be used).	
	Adverse effects: TESS <sup>2</sup> (only reported no adverse effects in electroacupuncture + antipsychotics group and compared ECG no difference between two groups).	
	Lab test: EEG; T <sub>3</sub> ; T <sub>4</sub> ; RUR; FT <sub>4</sub> I; TSH; rT <sub>3</sub> ; TG; TM; LH; FSH; T (not clinical outcomes).	
Notes	1. Two references with different participants' numbers but some data were the same and only extract- ed the same data with the same participants' number which could be used. No further detail.	
	2. The ratings were assessed before treatment, in 2nd, 4th, 6th and 8th week after treatment.	
	3. The effects were assessed after treatment with four degree which were worked out in Nanjing in 1958 (recovery, remarkable improvement, improvement and inefficacy).	

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "divided at random into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	High risk	Some of the graph showed data of 100 participants and some showed data of 69 participants. No further details.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	High risk	Some important data were the same but with different participants' number in two references. No further details.

### electro - Zhang 2001

Methods Alloc

Allocation: randomised.

Acupuncture for schizophrenia (Review)



### electro - Zhang 2001 (Continued)

	Blindness: not reporte Duration: 6 weeks.	d.	
Participants	Diagnosis: schizophrenia with depressive symptoms (ICD-10; CCMD-2-R).		
	N = 42. Age: between 17 and 5 Sex: 19 women and 23	4 years (mean age (32.19 ± 9.13) years). men.	
	History: duration of illr	ness between 7 months and 16 years (average (6.83 ± 4.77) years).	
	Setting: hospitalised p	atients.	
	Education level: highe	r than high school.	
	Country: China.		
	Inclusion criteria: had a CCMD-2-R and ICD-10 diagnosis of schizophrenia; HAMD ≥ 20 and without severe tendency of suicide; without severe somatic diseases; without extreme excited, stupor, silent and any other uncooperative state; all the patients received simple antipsychotics and without receiving any antidepressants and anxiolytics.		
Interventions	1. Electroacupuncture + antipsychotics: electroacupuncture (Intelligent electroacup and Yingtang; peak voltage 3-10 VP; for 45 minutes; once a day; 5 days a week except Sunday; 6 weeks as a treatment course) + antipsychotics (no further details). N = 22.		
	2. Antipsychotics: no fu	urther details. N = 20.	
Outcomes	Mental state: HAMD <sup>1</sup> , SDS.		
	Behaviour: leaving the study early.		
Notes	1. Another assessment standard (according to reduced rate): marked improvement (≥ 50%); improve- ment (≥ 25%); no effect (< 25%, reduced rate = [scores before treatment-scores after treatment]/scores before treatment*100%).		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly sampled".	
Allocation concealment (selection bias)	Unclear risk	Not stated.	
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.	
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.	
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.	
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.	

Acupuncture for schizophrenia (Review)



### electro - Zhang 2001 (Continued)

Other bias

Low risk

Not obvious.

#### electro - Zhou 1997

Methods	Allocation: randomised. Blindness: assessor blind. Duration: 6 weeks.		
Participants	Diagnosis: schizophrenia (DSM III; CCMD). N = 40. Age: mean age (31.8 ± 8.9) years. Sex: 17 women and 23 men.		
	History: average duratic Setting: hospitalised pa	on of illness (8.2 ± 7.0) years. tients.	
	Country: China.		
	Inclusion criteria: scores Exclusion criteria: sever	s higher than 35 points on BPRS <sup>1</sup> . e heart, liver, kidney or other somatic diseases.	
Interventions	1. Electroacupuncture + TCM types) + antipsycho	low dose antipsychotics: electroacupuncture <sup>*</sup> (acupoints choice according to $otics$ (a reduction of ~60% of their previously daily levels). N = 25.	
	2. Antipsychotics: contir ± 71.2) mg/day. N = 15.	nued to receive their usual antipsychotics; equivalent chlorpromazine dose (560	
	* Electroacupuncture: m tal acupoints (Zusanli [Y cycles/second; 500 ms p course (6 weeks).	nain acupoints (Yintang tou xinqu, Daling, Neiguan, Taiyang) and supplemen- 'ang deficiency], Sanyinjiao [Yin deficiency], Fenglong [ Persistent Phlegm]);180 pulse width; up to 60 mA; once a day except Sunday; 36 times as a treatment	
Outcomes	Global state: no clinicall Mental state: BPRS <sup>2,3</sup> .	y important change in global state <sup>1</sup> ; CGI (CGI-SI, CGI-GI, EI).	
	Behaviour: leaving the s Adverse effects: TESS.	tudy early.	
	Unable to use:		
	Lab test: cAMP; cGMP; th	ne neuropeptide beta-endorphin (not clinical outcome).	
Notes	1. Results were classified entation was recovered partly recovered); no eff	d as: marked improvement (the majority of symptoms were eliminated and ori- ); improvement (some of the symptoms were eliminated and orientation was fect (symptoms and orientation showed no change).	
	2.This rating scale was w and 6 weeks after treatm	vritten in Chinese and was assessed before treatment and at 2 weeks, 4 weeks nent.	
	3. Another assessment o ment (50%to 80%); sligh	riteria (according to reduced rate): marked improvement (≥ 80%); improve- nt improvement (20% to 50%); no effect (≤ 20%).	
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two study groups". No further details.	

Acupuncture for schizophrenia (Review)

### electro - Zhou 1997 (Continued)

Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Author compared two groups and did not use sham electroacupuncture.
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - "The subjects were evaluated clinically before and after the study course by specialists in psychiatry who were blinded as to which group the subjects belonged".
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### laser - Liu 1986

Methods	Allocation: randomised. Blindness: not reported. Duration: a treatment course.		
Participants	Diagnosis: schizophrenia with auditory hallucinations.		
	N = 60. Age: mean age (35.8 ± 13.0) years (He-Ne laser irradiation group (Ermen)); (33.2 ± 6.9) years (chlorpro- mazine + sham laser irradiation group); (37.7 ± 10.8) years (He-Ne laser irradiation group (non-acu- point)). Sex: 32 women and 28 men.		
	History: average duration of illness 5.1 years (He-Ne laser irradiation group (Ermen)); 6.2 years (chlor- promazine + sham laser irradiation group); 7.3 years (He-Ne laser irradiation group (non-acupoint)).		
	Country: China.		
	Inclusion criteria: had auditory hallucinations definitely schizophrenic in origin for more than one month.		
Interventions	1. He-Ne laser irradiation (Ermen): Ermen (double); irradiated with mouths partially open; the tube of the emitter held 30 cm away; power 4.7 hw; patch of red light 0.5 cm; wavelength 6328 Å; optimal cur- rent output 6 to 7.5 mA; for 15 minutes; once a day except Sunday; 30 times as a treatment course. N = 20.		
	2. Chlorpromazine + sham laser irradiation: chlorpromazine (average dosage 450 mg/d) + sham laser ir- radiation (the tube of the laser emitter pointed in the direction of the ear without real irradiation; once a day except Sunday; for 15 minutes; 30 times as a treatment course). N = 20.		
	3. He-Ne laser irradiation (non-acupoint): laser irradiation was thrown on the inner aspect of earlobe where no acupoint was located; once a day except Sunday; for 15 minutes; 30 times as a treatment course. N = 20.		
	Combination therapy: not received any supplementary medication.		

Acupuncture for schizophrenia (Review)


### laser - Liu 1986 (Continued)

 Outcomes
 Behaviour: leaving the study early.

 Adverse effects: numbness over the ear, upper extremity and chest on the treated side (only reported the data of the He-Ne laser irradiation group (Ermen)).

 Unable to use:

Mental state: rating scale of auditory hallucinations (for this particular trial).

Adverse effects: sensation of heat at irradiated site and a feeling of plugged auditory canal (equivocal data).

Notes

### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly numbered and treated as three groups".
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Author reported that three groups all used laser irradiation (two real and one sham) and one group combined with chlorpromazine but did not reported the other two groups used dummy drug.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### laser - Ma 1999

Methods	Allocation: randomised. Blindness: not reported. Duration: unclear.
Participants	Diagnosis: schizophrenia with auditory hallucinations (CCMD-2).
	N = 120. Age: between 17 and 53 years (mean age (32.02 ± 8.4) years) (He-Ne laser irradiation + chlorpromazine group); between 16 and 55 years (mean age (31.08 ± 8.6) years) (chlorpromazine group). Sex: 26 women and 94 men.
	History: duration of illness ≥ 15 days (He-Ne laser irradiation + chlorpromazine group); ≥ 10 days (chlor- promazine group).

Acupuncture for schizophrenia (Review)

laser - Ma 1999 (Continued)	Setting: hospitalised patients.			
	Country: China.			
	Inclusion criteria: had a CCMD-2 diagnosis of schizophrenia; BPRS ≥ 35.			
	Exclusion criteria: brain organic diseases or brain organic diseases.			
Interventions	1. He-Ne laser irradiation + chlorpromazine: He-Ne laser irradiation (Tinggong and Tinghui; wavelength 6328 Å; optimal current output $10 \pm mA$ ; negative ion generator $10^{-4}$ A; output total laser power 14-28 mµ; guided beam $\oplus 3 \times 300$ ; focus divergent optical lens; for 30 minutes; once a day except Sunday) + chlorpromazine (dose 300 to 550 mg/d; average dose (395 ± 55) mg/d). N = 60.			
	2. Chlorpromazine: dose 300 to 600 mg/d; average dose (400 $\pm$ 85 mg/d). N = 60.			
	Patients receiving antipsychotics before the study commenced needed a 2-week washout period.			
	Combination therapy: not combined with any other antipsychotics.			
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .			
	Mental state: BPRS (4 weeks); no clinically important change in specific symptoms (auditory hallucina- tions) <sup>2</sup> .			
	Behaviour: leaving the study early (4 week).			
	Service outcome: time to hospitalisation.			
	Others <sup>3</sup> : Time to auditory hallucinations disappeared.			
Notes	1. Assessment criteria: according to usual used 4 evaluation clinical criteria (recovery, marked improve- ment, improvement, no effect).			
	2. Assessment criteria: auditory hallucinations disappeared completely (disappeared completely); marked improvement (frequency of auditory hallucinations reduced obviously; clarity was significant- ly reduced; essentially no effect on patient's thinking or behaviour); improvement (frequency of audito- ry hallucinations reduced slightly; clarity was slightly reduced; partly effect on patient's thinking or be- haviour); no effect (no change of auditory hallucinations).			
	3. Author reported this outcome.			

Risk of bias
--------------

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Unclear risk	Sham laser irradiation not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias)	Low risk	No participants left the study early.

Acupuncture for schizophrenia (Review)



### **laser - Ma 1999** (Continued) Outcomes

Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### laser - Zhang 1991

Methods	Allocation: randomised. Blindness: assessor blind. Duration: 1 week washout period and 5 weeks treatment period.
Participants	Diagnosis: schizophrenia (paranoid type) (DSM III; CCMD-2-R).
	N = 31. Age: between 17 and 55 years (mean age 32.4 years). Sex: 11 women and 20 men.
	History: duration of illness between 3 months and 22 years (average duration (6.2 $\pm$ 5) years). Setting: hospitalised patients.
	Country: China.
	Exclusion criteria: somatic diseases.
Interventions	1. Laser acupuncture <sup>*</sup> : group 1 (Dazhui and Shenting); group 2 (Taiyang [double]). N = 11.
	2. Laser acupuncture + low dose chlorpromazine: laser acupuncture <sup>*</sup> (group 1 [Dazhui and Shenting] and group 2 [Taiyang [double]]) + chlorpromazine (150 to 300 mg/day). N = 10.
	3. Sham laser acupuncture + chlorpromazine: sham laser acupuncture (needles fixed with tape on acu- points) + chlorpromazine (350 to 600 mg/day). N = 10.
	*Laser acupuncture: using the two acupoints groups every other day alternately; needles inserted acu- points; for 15 minutes; once a day (except Sunday); 5 weeks as a treatment course; optical fibre output power > 2 MW, output laser distributing angle < 20°; core diameter of optical fibre 300 micron.
	Combination therapy (laser acupuncture group and laser acupuncture + low dose chlorpromazine group): only could use diazepam or chloral hydrate when patient with insomnia.
Outcomes	Global state: no clinically important change in global state <sup>1,2</sup> .
	Mental state: BPRS <sup>3</sup> .
	Behaviour: leaving the study early.
	Adverse effects: RESES <sup>3</sup> . Unable to use:
	Global state: CGI <sup>3</sup> (no useful data).
Notes	1. The rating was assessed after treatment.
	2. Assessment criteria (according to traditional criteria): recovery; marked improvement; improvement; no effect.
	3. The ratings were assessed before treatment and each week after treatment.
Risk of bias	

Acupuncture for schizophrenia (Review)

### laser - Zhang 1991 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "divided into laser acupuncture group, laser acupuncture and low dose chlorpromazine group and sham laser acupuncture and high dose chlorpromazine group using random allocation method". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Though author compared 3 groups and used sham laser acupuncture did not use dummy medications.
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - "two experience doctors assessed outcomes using blind evalu- ation method".
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### traditional - Bouhlel 2011 Methods Allocation: randomised. Blindness: double-blind. Duration: 24 days. Participants Diagnosis: schizophrenia or schizoaffective disorder (DSM IV). N = 36. Age<sup>1</sup>: between 21 and 56 years (mean age $(36.6 \pm 10.06)$ years). Sex:<sup>1</sup> 12 women and 19 men. History<sup>1</sup>: duration of disorders between 1 year and 37 years (average duration (12.68 ± 9.39) years) and the number of previous hospitalisations ranged from 0 to 28 times (average number $(6.19 \pm 7.02)$ times). Setting: hospitalised patients. Country: Tunisia. Inclusion criteria: patient who met DSM IV diagnosis of schizophrenia and who gave their consent. Exclusion criteria: patients who were suffering from organic disorder and those who were included in other research protocols. Interventions 1. Traditional acupuncture + antipsychotics: traditional acupuncture\* (local points, distal points diagnosed by TCM) + antipsychotics (no further details). $N = 15^{1}$ . 2. Sham acupuncture + antipsychotics: no further details. N = 16<sup>1</sup>. \* Traditional acupuncture: a simple manipulation; once for 20 minutes and 3 times a week; total 10 times.

Acupuncture for schizophrenia (Review)

traditional - Bouhlel 2011 (Continued)		
Outcomes	Mental state: PANSS; SAPS; SANS.	
	Unable to use:	
	Behaviour: leaving the study early (no useful data) (no further details).	
Notes	1. These data were from 31 participants.	
	2. The author reported outcomes of 31 hospitalised patients and reported that four other participants left the study before completing the 10 sessions and another patient left study after the first session, thus there were a total of 36 participants and we did not know which group the five participants who left the study early came from.	

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "a clinical randomised trial". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Low risk	Reports stated - "double-blind". The author reported that both the patient and the psychiatrist did not know if it was a true traditional acupuncture treatment or a placebo (sham acupuncture).
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Report stated - "outcome assessments were performed by the same treating psychiatrist and the psychiatrist did not know patient treated by a true tradi- tional acupuncture treatment or sham acupuncture".
Incomplete outcome data (attrition bias) Outcomes	High risk	Five participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### traditional - Liu 2010

Methods	Allocation: randomised. Blindness: not reported. Duration: 3 months.
Participants	Diagnosis: schizophrenia with refractory auditory hallucinations (CCMD-3). N = 100. Age: mean age (34.05 ± 8.35) years (traditional acupuncture + risperidone group); (35.12 ± 7.57) years (risperidone group). Sex: 43 women and 57 men. History: average duration of illness (4.27 ± 3.70) years (traditional acupuncture + risperidone group); (4.02 ± 3.91) years (risperidone group). Setting: hospitalised patients.

Acupuncture for schizophrenia (Review)

### traditional - Liu 2010 (Continued) Country: China.

	Inclusion criteria: had a CCMD-3 diagnosis of schizophrenia and agreed to receive treatment; hospi- talised and received systematic therapy with various antipsychotics, at least two, and the majority of psychiatric symptoms disappeared but the auditory hallucinations appeared repeatedly; had received other treatments (such as electric shock, acupuncture or Chinese herbs) but auditory hallucinations re- mained or reappeared; the duration of auditory hallucinations>8 weeks; BPRS < 30; expected length of hospitalisations > 3 months. Exclusion criteria: with allergic reactions or severe adverse effects; unwilling to receive treatment with the reasons of poor efficacy, side effects or others during trial process; with other severe physical com- plications during trial process.
Interventions	1. Traditional acupuncture + risperidone: traditional acupuncture <sup>*</sup> (used main acupoints <sup>**</sup> and adjunct acupoints <sup>***</sup> according to the TCM syndrome differentiation) + risperidone (Weisitong; initial dosage 1 mg [before breakfast]; gradually increased to the therapeutic dosage [2 to 6 mg/d] within 2 weeks; average dosage (2.3 ± 0.8) mg/d; daily total dosage less than 3 mg once a day before breakfast; otherwise twice a day before breakfast and dinner). N = 50.
	2. Risperidone: Weisitong; initial dosage 1 mg (before breakfast); gradually increased to the therapeutic dosage (2 to 6 mg/d) within 2 weeks; daily total dosage less than 3 mg once a day before breakfast; otherwise twice a day before breakfast and dinner. N = 50.
	combination therapy: could not combine with any other antipsychotics, antidepressants, mood sta- bilisers or electric shock; could add artane, scopolamine injections and propranolol with temporary therapeutic dosage according to the condition to enhance patient's compliance and stopped to use when patient's somatic complaints alleviated.
	* Traditional acupuncture: manipulated needles every 10 minutes; for 30 minutes; 4 to 5 times a week and no less than 4 times; 1 month as 1 treatment course.
	** Main acupoints: Shenmen (double); Daling (double); Taichong (double); Tinggong(double); Yifeng (double); Baihui.
	*** Adjunct acupoints:
	Type of stagnation of phlegm and Qi: Fenglong (double) and Tanzhong (double).
	Type of failure of the heart and kidney integrating: Taixi (double) and Laogong (double).
	Type of phlegm-fire attacking upwards: Yongquan (double) and Houxi (double)
	Type of deficiency of both the heart and spleen: Zusanli (double) and Sanyinjiao (double).
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .
	Mental state <sup>2</sup> : BPRS; SAHS.
	Behaviour: leaving the study early.
	Adverse effects: TESS <sup>3</sup> .
Notes	1. Assessment criteria: recovery (auditory hallucinations disappeared completely; without any other psychiatric symptoms; insight recovered completely); marked improvement (the majority of auditory hallucinations disappeared; other psychiatric symptoms improved; insight partly recovered); improvement (the number of auditory hallucinations reduced; voice clarity was fuzzy; partly affected patient's daily life; insight was not complete); no effect (the number, duration and content of auditory hallucinations without marked change or became worse).
	2. The ratings were assessed before treatment and at the end of 1, 2 and 3 treatment courses.
	3. The rating was assessed at the end of 1, 2 and 3 treatment courses but lab test repeated twice (be- fore and after third treatment).

Acupuncture for schizophrenia (Review)

### traditional - Liu 2010 (Continued)

### **Risk of bias**

Bias

Cochrane Database of Systematic Reviews

Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly into two groups using the random number table ac- cording to the admission order".
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	High risk	Four participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Unclear risk	This study was the Lanzhou's 2008 second batch technology program.

traditional - Luo 2006	
Methods	Allocation: randomised. Blindness: single-blind <sup>1</sup> . Duration: 1 week washout period and 3 months treatment courses.
Participants	Diagnosis: type II syndrome of schizophrenia (CCMD-3). N = 60. Age: between 15 and 54 years (mean age (26.81 ± 7.43) years) (traditional acupuncture + risperidone group); between 16 and 56 years (mean age (27.47 ± 6.78) years) (risperidone group). Sex: 26 women and 34 men.
	History: duration of illness between 1.5 and 18 years (average duration (6.81 ± 5.42) years) (tradition- al acupuncture + risperidone group); between 1.5 and 20 years (average duration (7.73 ± 5.18) years) (risperidone group).
	Country: China.
	Inclusion criteria: had a CCMD-3 diagnosis of schizophrenia; BPRS > 35; SANS $\ge$ 60; SAPS < 8.
	Exclusion criteria: severe somatic diseases; cerebral organic psychosis; ethyl alcohol and drug depen- dence.
Interventions	1. Traditional acupuncture + risperidone: traditional acupuncture <sup>*</sup> (two acupoints groups <sup>**</sup> in turn) + risperidone (initial dose 1 mg/d; added to 4 to 6 mg/d within 10 days; twice a day). N = 30.
	2.Risperidone: initial dose 1 mg/d; added to 4 to 6 mg/d within 10 days; twice a day. N = 30.

Acupuncture for schizophrenia (Review)



Risk of bias	
	3. The ratings were assessed before treatment and at 1 month, 2 moths and 3 months after treatment.
	2. Assessment criteria (according to criteria of Neuropsychiatric Association of Chinese Medical Associ- ation): recovery; marked improvement; effect; no effect.
Notes	1. Contact made with author Cheng Luo, however, still unclear about single-blind method.
	Behaviour: leaving the study early.
	Mental state <sup>3</sup> : BPRS; SANS.
Outcomes	Global state: no clinically important change in global state <sup>2</sup> .
	Group B: Sishencong (reinforcing method); Yingtang Tou face acupuncture heart area (reinforcing method); Zusanli (right), Yanglingquan (right), Neiguan (right), Shenmen (right), Yongquan (right) (rein- forcing-reducing method).
	Group A: Baihui (reinforcing method); Shenting Tou Shangxing (reinforcing method); Zusanli (left), Yan- glingquan (left), Neiguan (left), Shenmen (left), Yongquan (left) (reinforcing-reducing method).
	** Acupoints group:
	* Traditinal acupuncture: for 30 minutes; once a day; 20 days as a treatment course and began another treatment course after 10 non-treatment days.
	Combination therapy: supporting symptomatic treatment, psychological and occupational and recre- ational rehabilitation.
traditional - Luo 2006 (Continu	ed)

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	Unclear risk	Report stated - "using single-blind method". No further details.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Report stated - "using single-blind method". No further details.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

traditional - Ma 2008			
Methods	Allocation: randomised Blindness: not reported Duration: 1 week wash	l. J. out period and 6 weeks treatment period.	
Participants	Diagnosis: schizophrenia (type of stagnation of phlegm and Qi) (CCMD-3, diagnosis of TCM).		
	Age: between 16 and 48 group); between 17 and Sex: 22 women and 38	8 years (mean age (29.2 ± 6.2) years) (traditional acupuncture + risperidone d 51 years (mean age (30.6 ± 3.6) years) (risperidone group). men.	
	History: duration of illn tional acupuncture + ri: years) (risperidone grou Setting: both inpatient:	ess between 1 month and 9 years (average duration (4.6 ± 3.42) years) (tradi- speridone group); between 2 months and 11 years (average duration (5.3 ± 3.6) up). s and outpatients.	
	Country: China.		
	Inclusion criteria: had a stagnation of phlegm a	a CCMD-3 diagnosis of schizophrenia; PANSS ≥ 60; had a TCM diagnosis of type of nd Qi.	
	Exclusion criteria: preg dence.	nant or lactating women; severe organic diseases; alcohol and drug depen-	
Interventions 1. Traditional acupuncture + risperidone: tradit shi, Fenglong, Daling, Yingtang) + risperidone (S weeks). N = 30.		ture + risperidone: traditional acupuncture <sup>*</sup> (Juque, Tanzhong, Taichong, Jian- 'ingtang) + risperidone (Silishu; initial dose 1 mg/d; added to 2 to 4 mg/d within 2	
	2.Risperidone: Silishu; initial dose 1 mg/d; added to 2 to 6 mg/d within 2 weeks. N = 30.		
	* Traditional acupuncto with 2 days non-treatm	ure: reducing by rotating needles; for 30 minutes; once a day; 5 days treatment nent interval.	
Outcomes	Mental state: PANSS <sup>1,2</sup> .		
	Behaviour: leaving the	study early.	
	Adverse effects: TESS <sup>1</sup> .		
	Unable to use:		
	Adverse effects: lab tes	t (kidney function test -not reported.	
Notes	1. The ratings were assessed before treatment and at 2 weeks, 4 weeks and 6 weeks after treatment.		
	2. Another assessment standard (according to reduced score): recovery (≥ 75%); marked improvement (50% to 74%); improvement (25% to 49%); no effect (< 25%).		
Risk of bias			
Bias	Authors' judgement	Support for judgement	
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.	
Allocation concealment (selection bias)	Unclear risk	Not stated.	
Blinding of participants and personnel (perfor- mance bias)	High risk	Sham electroacupuncture not used.	

Acupuncture for schizophrenia (Review)



### traditional - Ma 2008 (Continued) All outcomes

Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	<ol> <li>We were unable to locate the protocol, unsure if selective reporting oc- curred.</li> <li>Kidney function tests were completed but not reported and we are unsure of the reason for this omission.</li> </ol>
Other bias	Low risk	Not obvious.

Allocation: randomised. Blindness: not reported. Duration: 3 months.		
Diagnosis: schizophrenia with auditory hallucinations (CCMD-3). N = 84. Sex: 20 women and 64 men. Setting: hospitalised patients. Country: China. Inclusion criteria: had a CCMD-3 diagnosis of schizophrenia; duration of hospitalisation>6 months; re-		
ceived antipsychotics with good compliance but auditory hallucinations did not disappear.		
<ol> <li>Traditional acupuncture + antipsychotics: traditional acupuncture<sup>*</sup> (acupoints choice according to the type of TCM<sup>**</sup>) + antipsychotics (remained previous antipsychotics treatment). N = 42.</li> </ol>		
2. Antipsychotics: remained previous antipsychotics treatment. N = 42.		
Combination therapy: could add artane, propranolol and benzodiazepine drugs and could not received any other antipsychotics.		
* Traditional acupuncture: acupoint near ear and on the head used transport point needling method and twisting and reducing method; acupoint on the limbs used quick-slow supplementation and drain- ing method and directional supplementation and draining method; manipulated needles every 10 min- utes; for 30 minutes; 3 to 4 times a week and not less than 3 times; 3 weeks as a treatment course and 1- week interval between two treatment courses; total 3 treatment courses.		
** Acupoints choice according to type of TCM:		
Type of stagnation of phlegm and Qi: Taichong, Fenglong, Tinggong, Yifeng, Baihui, Daling, Tanzhong.		
Type of failure of the heart and kidney integrating: Tinggong, Taixi, Shenmen, Sanyingjiao, Baihui, Tongtian.		
Type of phlegm-fire attacking upwards: Laogong, Yongquan, Daling, Taixi, Yifeng, Shenmen, Quchi.		
Type of deficiency of both the heart and spleen: Baihui, Pishu, Xinshu, Shenshu, Sanyingjiao, Zusanli.		

Acupuncture for schizophrenia (Review)

### traditional - Tang 2005 (Continued)

Outcomes	Global state: no clinically important change in global state <sup>1</sup> .
	Behaviour: leaving the study early.
Notes	1. Assessment criteria: marked improvement (auditory hallucinations disappeared; psychiatric symp- toms cause by auditory hallucinations improved markedly); improvement (times of auditory hallucina- tions reduced or sound was fuzzy; other psychiatric symptoms improved); no effect (auditory halluci- nations and psychiatric symptoms did not improve).

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided into two groups". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

### traditional - Wang 2006 Methods Allocation: randomised. Blindness: not reported. Duration: 30 days. Participants Diagnosis: hebephrenic schizophrenia (CCMD-2-R). N = 48. Age: mean age $(15.87 \pm 2.48)$ years (traditional acupuncture + low dose antipsychotics group); $(16.11 \pm 10^{-1})$ 2.73 years (traditional acupuncture group); (15.74 ± 2.89) years (traditional antipsychotics group). Sex: 18 women and 30 men. History: average duration of illness (1.54 ± 1.29) years (traditional acupuncture + low dose antipsychotics group); (1.60 ± 1.21) years (traditional acupuncture group); (1.52 ± 1.17) years (traditional antipsychotics group). Setting: hospitalised patients. Country: China.

Acupuncture for schizophrenia (Review)

traditional - Wang 2006 (Cont	<sup>inued)</sup> Inclusion criteria: had a	a CCMD-2-R diagnosis of hebephrenic schizophrenia; BPRS ≥ 35.		
	Exclusion criteria: had ent treatments caused ing the study period; cl	a CCMD-2-R diagnosis of hebephrenic schizophrenia but BPRS < 35; with differ- by combining with other diseases; could not continue to receive treatment dur- nanged to other treatments.		
Interventions	<ol> <li>Traditional acupuncture + low dose antipsychotics: traditional acupuncture<sup>*</sup> (Taichong [reducing method], Hegu [reducing method], Neiguan [straight inserted], Daling [straight inserted], Renzhong [re- ducing method], Dazhui [straight inserted] + acupoints choice according to type of TCM<sup>**</sup>) + low dose antipsychotics (≤equivalent chlorpromazine dose 0.2 g/d). N = 16.</li> </ol>			
	2. Traditional acupuncture*:Taichong (reducing method), Hegu (reducing method), Neiguan (straight inserted), Daling (straight inserted), Renzhong (reducing method), Dazhui (straight inserted) + acupoints choice according to type of TCM**. N = 16.			
	3. Antipsychotics: enou	3. Antipsychotics: enough dose antipsychotics (equivalent chlorpromazine dose 0.4 to 0.6 g/d). N = 16.		
	Combination therapy:	psychological counsel every day with parents according to patient's condition.		
	* Traditional acupunct minutes; once a day; 1	ure: continued to manipulate needles 3 to 5 minutes every ten minutes; for 45 5 times as a treatment course; 2 treatment courses.		
	** Acupoints choice ac	cording to type of TCM:		
	Type of Qi stagnation and blood stasis: Yanglingquan, Sanyingjiao.			
	Type of Internal retention of phlegm and dampness; Fenglong.			
	Type of internal disturbance of pyrophlegm: Fenglong, Xingjian.			
	Type of Yin deficiency and fire excess: Fuliu.			
	Type of Yang deficiency: Shenshu (moxibustion), Pishu, Guanyuan (moxibustion).			
Outcomes	Global state: no clinically important change in global state <sup>1,2</sup> .			
	Mental state <sup>3</sup> : BPRS, SAPS, SANS.			
	Behaviour: leaving the study early.			
	Adverse effects <sup>3</sup> : TESS.			
Notes	1. Assessment criteria: recovery (psychiatric symptoms disappeared completely; insight recovered); ef- fect (marked improvement [psychiatric symptoms disappeared completely or the majority main psy- chiatric symptoms disappeared; insight partly recovered or not recovered]; improvement [symptoms disappeared partly; insight did not recover); no effect (no symptoms improved obviously and insight did not recover).			
	2. The rating was assessed 30 days after treatment.			
	3. The ratings were assessed before treatment and each 15 days after treatment.			
Risk of bias				
Bias	Authors' judgement	Support for judgement		
Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly divided into 3 groups using random allocation table according to the admission date and case number".		
Allocation concealment (selection bias)	Unclear risk	Not stated.		

Acupuncture for schizophrenia (Review)

### traditional - Wang 2006 (Continued)

Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Sham electroacupuncture not used.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Low risk	Not obvious.

traditional - Xu 2004	
Methods	Allocation: randomised. Blindness: not reported. Duration: 80 days.
Participants	Diagnosis: schizophrenia (CCMD-2-R). N = 80.
	Age: mean age (31.74 ± 10.23) years (traditional acupuncture + low dose antipsychotics group); (30.12 ± 8.06) years (antipsychotics group). Sex: 80 men and no women.
	History: average duration of illness (5.53 ± 6.08) years (start age (25.33 ± 8.12) years) (traditional acupuncture + low dose antipsychotics group); (6.04 ± 3.47) years (start age (26.17 ± 7.32) years) (an- tipsychotics group).
	Type: hebephrenic schizophrenia 2 cases, paranoid type 28 cases, simple schizophrenia 1 case, undif- ferentiated type 9 cases (traditional acupuncture + low dose antipsychotics group); hebephrenic schiz- ophrenia 1 case, paranoid type 26 cases, simple schizophrenia 2 cases, catatonic type 1 case, undiffer- entiated type 10 cases (antipsychotics group) Setting: hospitalised patients.
	Country: China.
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; BPRS ≥ 35.
	Exclusion criteria: had a CCMD-2-R diagnosis of schizophrenia but BPRS < 35 or had diseases which would affect trial results.
Interventions	<ol> <li>Traditional acupuncture + low dose antipsychotics: traditional acupuncture<sup>*</sup> (used main acupoints<sup>**</sup> and adjunct acupoints<sup>***</sup> [according to the TCM type of schizophrenia] and special acupoints<sup>****</sup> [ac- cording to symptoms]) + low dose antipsychotics (equivalent chlorpromazine dose ≤ 0.2 g/d). N = 40.</li> </ol>
	2. Antipsychotics: maximum daily dose equivalent chlorpromazine dose 0.4 to 0.7 g/d. N = 40.
	Combination therapy: added artane when patient with severe extrapyramidal side effects; added pro- pranolol when patient with severe palpitation and heart rate > 100 times/minute.
	* Traditional acupuncture: used lifting-thrusting supplementation and draining method and twirling supplementation and draining method according to patient's condition; manipulated needles about

Acupuncture for schizophrenia (Review)



traditional - Xu 2004 (Continued	ed) 3 minutes and once ten minutes; total for about 30 minutes; at first once a day then reduced to once every two days when patient was in stable condition and once a week after psychiatric symptoms dis- appeared.				
	** Main acupoints: Shuigou, Baihui, Neiguan, Sanyingjiao.				
	***Adjunct acupoints (according to the CTM type of schizophrenia):				
	Type of internal disturbance of pyrophlegm: Zhongwan, Fenglong, Xingjian.				
	Type of Internal Retention of phlegm and dampness: Fenglong, Yinglingquan, Zusanli.				
	Type of Qi stagnation and Blood stasis: Xuehai, Geshu.				
	Type of Yin deficiency and fire excess: Shenmen, Fuliu.				
	Type of Yang deficiency	r: Taixi, Guanyuan (moxibustion).			
	Other miscellaneous ty	pes such as deficiency of both heart and spleen: Anmian, Shenmen.			
_	**** Special acupoints (according to symptoms): Zhongzhu, Tinggong (auditory hallucinations); Yuyao (phosphenes); Hegu, Laogong (agitation).				
Outcomes	Mental state <sup>1</sup> : BPRS; SAPS; SANS.				
	Behaviour: leaving the	study early.			
	Adverse effects: TESS <sup>2</sup> .				
	Unable to use: Global state: no clinically important change in global state <sup>3,4</sup> (only reported P value, P value > 0.05).				
Notes	1. The ratings were assessed before treatment and each 20 days after treatment. 2. The rating was assessed each 20 days after treatment.				
	3. Assessment criteria: recovery (psychiatric symptoms disappeared completely; insight recovered); marked improvement (psychiatric symptoms disappeared completely or the majority main psychiatric symptoms disappeared; insight partly recovered or not recovered); improvement (symptoms disap- peared partly; insight did not recover); no effect (no symptoms improved obviously and insight did not recover).				
	4. The rating was asses	sed after treatment.			
Risk of bias					
Bias	Authors' judgement	Support for judgement			
Random sequence genera- tion (selection bias)	Low risk	Report stated - "randomly divided into experimental group and control group using random allocation table according to the admission date and case number".			
Allocation concealment (selection bias)	Unclear risk	Not stated.			
Blinding of participants and personnel (perfor-	High risk     Sham electroacupuncture not used.				

All outcomes Blinding of outcome assessment (detection bias) All outcomes

Acupuncture for schizophrenia (Review)

mance bias)

\_

### traditional - Xu 2004 (Continued)

Incomplete outcome data (attrition bias) Outcomes	Low risk	Three participants left the study early but results graphs (BPRS, SAPS, SANS, TESS) showed that author analysed 40 cases in each group.
Selective reporting (re- porting bias)	Unclear risk	We were unable to locate the protocol, unsure if selective reporting occurred.
Other bias	Unclear risk	Participants were men and unsure if gender selection bias occurred.

### traditional - Zhao 2005a

Methods	Allocation: randomised. Blindness: not reported. Duration: 60 days.
Participants	Diagnosis: schizophrenia (type I; Kuang type) (CCMD-2-R; Diagnosis and Effect Standard of Diseases in TCM [1994]) . N = 300.
	Sex: 162 women and 138 men.
	History: longer than 3 months.
	Setting: outpatients.
	Country: China.
	Inclusion criteria: had a CCMD-2-R diagnosis of schizophrenia; had a TCM diagnosis of Kuang type of phlegm-fire disturbing mind; stopped to receive relative drug treatment more than 1 month.
	Exclusion criteria: cerebral organic mental disorders; mental disorders caused by somatic diseases; mental disorders caused by psychoactive substances and non-dependent substances; had both schizo- phrenia and mood disorders symptoms but the duration of illness of disruptive symptoms was shorter than that of mood disorder symptoms or no longer than 2 weeks; with severe cardiovascular and cere- brovascular, liver and haematopoietic system diseases.
Interventions	1. Fuyuankang capsule: 5 capsules (0.4 g/capsule) every time; 3 times a day; with warm water. N = 90.
	2. Traditional acupuncture: Shuigou, Shaoshang, Yingbai, Fengfu, Daling, Quchi, Fenglong; once a day; for 30 minutes. N = 90.
	3. Traditional acupuncture + Fuyankang capsule: traditional acupuncture (Shuigou, Shaoshang, Ying- bai, Fengfu, Daling, Quchi, Fenglong; once a day; for 30 minutes) + Fuyuankang capsule (5 capsules every time; 3 times a day; with warm water). N = 90.
	4. Weisitong (risperidone tablet): initial dose 1 mg/d (after dinner); added 1 mg after 3 days then added 0.1 mg every week until daily dose 4 mg. N = 30.
	* Author used placebo with therapeutic medications; no further details.
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .
	Behaviour: leaving the study early.
	Unable to use:
	Global state: TCM Syndromes Scale <sup>2</sup> (no useful data).
	Mental state: BPRS (no data).

Acupuncture for schizophrenia (Review)

#### traditional - Zhao 2005a (Continued)

Adverse effects: lab test (ECG; blood routine test; urine routine test; stool test; liver function; kidney function); blood pressure; adverse reaction (only reported data for traditional acupuncture + Fuyuankang group).

Lab test: SOD (not clinical outcomes).

Notes1. Assessment criteria: recovery (talk, behaviour and expression all returned to normal state; could deal<br/>with daily affairs; the reduction of BPRS ≥ 95%; improvement (talk, behaviour and expression returned<br/>to normal state approximately or obviously improved; the reduction of BPRS 30% to 94%); no effect<br/>(talk, behaviour and expression still existed as before; the reduction of BPRS < 29%).</th>

2. Author also reported the global state of TCM.

3. The study of 'Zhao 2005' consisted of two parts - 'traditional - Zhao 2005a' (type I schizophrenia) and 'traditional - Zhao 2005b' (type II schizophrenia).

4. The allocation rate of groups of each type schizophrenia was 3:3:3:1.

### **Risk of bias**

Bias	Authors' judgement	Support for judgement
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided". No further details.
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Used placebo but did not use sham acupuncture.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	High risk	Although the protocol was not located, the author reported that BPRS was one of the outcomes, however, only reported global state.
Other bias	Low risk	Not obvious.

### traditional - Zhao 2005b

Methods	Allocation: randomised. Blindness: not reported. Duration: 60 days.
Participants	Diagnosis: schizophrenia (type II; Dian type) (CCMD-2-R; Diagnosis and Effect Standard of Diseases in TCM [1994]). N = 300. Sex: 156 women and 144 men. History: longer than 3 months.

Acupuncture for schizophrenia (Review)

traditional - Zhao 2005b (Con	<sup>tinued)</sup> Setting: outpatients.				
	Country: China.				
	Inclusion criteria: had a phlegm-Qi stagnation a	CCMD-2-R diagnosis of schizophrenia; had a TCM diagnosis of Dian type of and blinding; stopped to receive relative drug treatment more than 1 month.			
	Exclusion criteria: cerel mental disorders cause phrenia and mood diso than that of mood diso brovascular, liver and h	oral organic mental disorders; mental disorders caused by somatic diseases; d by psychoactive substances and non-dependent substances; had both schizo- rders symptoms but the duration of illness of disruptive symptoms was shorter rder symptoms or no longer than 2 weeks; with severe cardiovascular and cere- aematopoietic system diseases.			
Interventions	1. Fuyuankang capsule:	5 capsules (0.4 g/capsule) every time; 3 times a day; with warm water. N = 90.			
	2. Traditional acupunct 90.	ure: Xinshu, Ganshu, Pishu, Shenmen, Fenglong; once a day; for 30 minutes. N =			
	3. Traditional acupunct Shenmen, Fenglong; or a day; with warm water	cure + Fuyankang capsule: traditional acupuncture (Xinshu, Ganshu, Pishu, nce a day; for 30 minutes) + Fuyuankang capsule (5 capsules every time; 3 times ). N = 90.			
	4. Weisitong (risperidor 0.1 mg every week unti	ne tablet): initial dose 1 mg/d (after dinner); added 1 mg after 3 days then added I daily dose 4 mg. N = 30.			
	* Author used placebo	with therapeutic medications; no further details.			
Outcomes	Global state: no clinically important change in global state <sup>1</sup> .				
	Behaviour: leaving the study early.				
	Unable to use:				
	Global state: TCM Synd	romes Scale <sup>2</sup> (no useful data).			
	Mental state: BPRS.				
	Adverse effects: lab test ney function); blood pro Fuyuankang group).	t (ECG; blood routine test; urine routine test; stool test; liver function; kid- essure; adverse reaction (only reported data of traditional acupuncture +			
	Lab test: SOD (not clinio	cal outcomes).			
Notes	<ol> <li>Criteria: recovery (talk, behaviour and expression all returned to normal state; could deal v affairs; the reduction of BPRS ≥ 95%; improvement (talk, behaviour and expression returned state approximately or obviously improved; the reduction of BPRS 30% to 94%); no effect (ta iour and expression still existed as before; the reduction of BPRS &lt; 29%).</li> </ol>				
	2. Author also reported	the global state of TCM.			
	3. The study of 'Zhao 2005' consisted of two part - 'traditional - Zhao 2005a' (type I schizophrenia) and 'traditional - Zhao 2005b' (type II schizophrenia).				
	4. The allocation rate of groups of each type schizophrenia was 3:3:3:1.				
Risk of bias					
Bias	Authors' judgement	Support for judgement			
Random sequence genera- tion (selection bias)	Unclear risk	Report stated - "randomly divided". No further details.			

Acupuncture for schizophrenia (Review)



### traditional - Zhao 2005b (Continued)

	nued)	
Allocation concealment (selection bias)	Unclear risk	Not stated.
Blinding of participants and personnel (perfor- mance bias) All outcomes	High risk	Used placebo but did not use sham acupuncture.
Blinding of outcome as- sessment (detection bias) All outcomes	Unclear risk	Not stated.
Incomplete outcome data (attrition bias) Outcomes	Low risk	No participants left the study early.
Selective reporting (re- porting bias)	High risk	Although the protocol was not located, the author reported that BPRS was one of the outcomes, however, only reported global state.
Other bias	Low risk	Not obvious.
DSM: Diagnostic and Statistical ICD 10: International Classificat CCMD: Chinese Classification o Global state: CGI: Clinical Global Impression CGI-SI: Clinical Global Impression CGI-GI: Clinical Global Impression CGI-EI: Clinical Global Impression Mental state: BPRS: Brief Psychiatric Rating S PANSS: Positive and Negative S SANS: Scale for the Assessment SAPS: Cug Self-Rating Depressi PSYRATS-AH: Psychotic Sympto Adverse effects: TESS: Treatment Emergent Syn RESES: Rating Scale for Extrapy Test: ECG: electrocardiogram. EEG: electrocardiogram. CAMP: cyclic guanosine monop CGMP: cyclic guanosine monop CGMP: cyclic guanosine monop CGMP: cyclic denosine denosine CGMP: cyclic denosine denosine CGMP: cyclic	Manual. tion of Diseases. f Mental Disorders. Scale. on - Severity of illness. on - Improvement scale. on - Efficacy Index. Scale. Syndrome Scale. t of Negative Symptoms. ination Scale. or Depression. on Scale. om Rating Scales Auditor nptom Scale. matinal Side Effects.	y Hallucination Subscale.

T: testosterone.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



SOD: superoxide dismutase. Others: TCM: Traditional Chinese Medicine. CCEA: Computer-Controlled Electric Acupuncture. mg/d: mg per day.

### Characteristics of excluded studies [ordered by study ID]

Study	Reason for exclusion
Luo 2006a	Allocation: randomised.
	Participants: people with schizophrenia or mood disorders (affective disorders) manic episode.
	Interventions: acupuncture + antipsychotics vs antipsychotics.
	Outcomes: not reported by diagnostic group.
Ma 2002	Allocation: randomised.
	Participants: people with chronic schizophrenia.
	Interventions: antipsychotics + electroacupuncture (Yingtang and Baihui) versus antipsychotics + electroacupuncture (Baihui and Spiritual and emotional movement area) (compared elec- troacupuncture with different acupoints), not acupuncture vs other treatments.
Ma 2004	Allocation: not randomised.
Sun 1994	Allocation: not randomised, case series.
Wu 2004	Allocation: randomised.
	Participants: people with schizophrenia.
	Interventions: acupoint injection + low dose chlorpromazine (Wintermin) versus chlorpromazine (Wintermin).
	Outcomes: BPRS (mental state); TESS (adverse effects).
	* Some important data from this study (conducted in same institution as acupoint inj - Pan 2002) were the same as acupoint inj - Pan 2002 (baseline and endpoint total BPRS), but numbers allocated, the acupoint method, and BPRS' factor data were different; we could not contact author of Wu 2004 but author of acupoint inj - Pan 2002 had no knowledge of this study.
Xiong 2009	Allocation: not randomised, quasi-randomised (according to the admission order; patient with odd number into electroacupuncture group and with even number into MECT group).
Xue 1985	Allocation: divided randomly to three intervention groups but unclear how allocated within one key intervention group
	Participants: people with schizophrenia.
	Interventions: Electroacupuncture/acuclip convulsive therapy (EACT) (involved mixture of tech- niques - not all of which are acupuncture - several of which could be applied in a single person) ver- sus electroconvulsive therapy (ECT) versus ECT or EACT (unclear how allocation to EACT undertak- en within this group), not acupuncture versus none/other treatment.
Zhong 1995	Allocation: randomised. Participants: people with common mental diseases. Interventions: electroacupuncture versus computerised electrode.

Acupuncture for schizophrenia (Review)



### Study

**Reason for exclusion** 

Zhuge 1993

Allocation: not randomised, allocated according to age and sex.

Mental state: BPRS: Brief Psychiatric Rating Scale. Adverse effects: TESS: Treatment Emergent Symptom Scale.

Test: ECT: electroconvulsive therapy. EACT: electric acupuncture convulsive therapy. Other: MECT: modified electroconvulsive therapy

### Characteristics of studies awaiting assessment [ordered by study ID]

### NCT01167348

Methods	Allocation: randomised. Blindness: single-blind (assessor). Duration: 2 months. Design: parallel.
Participants	Diagnosis: schizophrenia with obesity (ICD).
	N = 86 (estimated enrolment).
	Sex: women and men.
	Inclusion criteria: ICD:295, living in chronic ward for more than 2 months; body mass index (BMI) > or = 24; aged between 20 to 60; psychotic status stable and can communicate.
	Exclusion criteria: unstable psychotic status; participants who have endocrine disease; participants who have cardiac disease; participants who have immunological disease; participants who have liver or renal function impairment; pregnant or lactating woman; cerebrovascular accident (CVA) stroke and disability; participants who attend weight control programs in last 3 months; any conditions that treating doctors refuse to join in this study.
Interventions	1. Auricular acupressure: with a 0.5 mm x 0.5 mm seed on the specific positions on the ear and press these specific positions 3 times a day.
	2. Placebo: no further details.
Outcomes	Metabolic: Body weight scale, waste circumference, fat percentage by BIA.
Notes	Contacted with Han-Yi Ching using the email address from the ClinicalTrials.gov but no feedback. No further details and unable to confirm whether the study has finished and unable to locate the full text.

BIA: Bio-Impedance Analysis

ICD: International Classification of Diseases.

### DATA AND ANALYSES

## Comparison 1. ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Global state: Not improved, endpoint	4		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
1.1 medium-term (various similar criteria)	3	244	Risk Ratio (M-H, Fixed, 95% CI)	0.40 [0.28, 0.57]
1.2 duration unclear	1	120	Risk Ratio (M-H, Fixed, 95% CI)	0.2 [0.01, 4.08]
2 Mental state: 1a. General - average score (BPRS, endpoint, high score = worse)	5		Mean Difference (IV, Fixed, 95% CI)	Subtotals only
2.1 short-term	4	327	Mean Difference (IV, Fixed, 95% CI)	-4.32 [-5.28, -3.36]
2.2 medium-term	3	220	Mean Difference (IV, Fixed, 95% CI)	-5.51 [-6.71, -4.30]
3 Mental state: 1b. General - average score (BPRS, endpoint, high score = worse, medium-term) - subgroup analysis	3		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
3.1 traditional acupuncture	2	156	Mean Difference (IV, Ran- dom, 95% CI)	-7.95 [-14.29, -1.61]
3.2 acupoint injection	1	64	Mean Difference (IV, Ran- dom, 95% CI)	-0.70 [-5.02, 3.62]
4 Mental state: 2a. General - average score (PANSS, endpoint, high score = worse)	4		Mean Difference (IV, Fixed, 95% CI)	Subtotals only
4.1 short-term	4	245	Mean Difference (IV, Fixed, 95% CI)	-2.47 [-4.31, -0.63]
4.2 medium-term	2	135	Mean Difference (IV, Fixed, 95% CI)	-3.79 [-6.43, -1.15]
5 Mental state: 2b. General - average score (PANSS, endpoint, high score = worse, short-term) - subgroup analysis	4		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
5.1 traditional acupuncture	1	60	Mean Difference (IV, Ran- dom, 95% CI)	1.30 [-1.56, 4.16]
5.2 electroacupuncture	3	185	Mean Difference (IV, Ran- dom, 95% CI)	-5.64 [-10.93, -0.35]
6 Mental state: 2c. General - average score (PANSS, endpoint, high score = worse, short-term) - electroacupuncture sub- group analysis	2	135	Mean Difference (IV, Fixed, 95% CI)	-3.79 [-6.43, -1.15]

Acupuncture for schizophrenia (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
7 Mental state: 2d. General - average score (PANSS, endpoint, high score = worse) - Skewed data			Other data	No numeric data
7.1 short-term			Other data	No numeric data
7.2 medium-term			Other data	No numeric data
8 Mental state: 2e. General - not improved (PANSS), endpoint	4		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
8.1 reduced rate < 25%, short-term	3	197	Risk Ratio (M-H, Fixed, 95% CI)	0.65 [0.45, 0.94]
8.2 reduced rate < 30%, short-term	1	90	Risk Ratio (M-H, Fixed, 95% CI)	0.91 [0.43, 1.92]
8.3 reduced rate < 30%, follow-up, medi- um-term	1	90	Risk Ratio (M-H, Fixed, 95% Cl)	0.69 [0.36, 1.31]
9 Mental state: 3. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data			Other data	No numeric data
10 Mental state: 4a.Specific - average score - negative symptoms (SANS, end- point, high score = worse)	1		Mean Difference (IV, Fixed, 95% CI)	Subtotals only
10.1 short-term	1	60	Mean Difference (IV, Fixed, 95% CI)	-7.66 [-13.05, -2.27]
10.2 medium-term	1	60	Mean Difference (IV, Fixed, 95% CI)	-12.35 [-17.54, -7.16]
11 Mental state: 4b. Specific - average score - negative symptoms (SANS, end- point, high score = worse, short-term) - Skewed data			Other data	No numeric data
11.1 traditional acupuncture			Other data	No numeric data
11.2 electroacupuncture			Other data	No numeric data
12 Mental state: 5a. Specific - average score - depression (HAMD, endpoint, high score = worse, short-term)	2	109	Mean Difference (IV, Ran- dom, 95% CI)	-8.66 [-12.10, -5.22]
13 Mental state: 5b. Specific - not im- proved - depression (HAMD, reduced rate < 25%, short-term)	2	109	Risk Ratio (M-H, Fixed, 95% CI)	0.17 [0.08, 0.34]
14 Mental state: 7b. Specific - not im- proved - auditory hallucinations (PSYRAS- AH, reduction < 20%, short-term)	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.27 [0.14, 0.52]

Acupuncture for schizophrenia (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
15 Mental state: 7a. Specific - average score - auditory hallucinations (PSYRAS- AH,endpoint, high score = worse, short- term)	1	60	Mean Difference (IV, Fixed, 95% CI)	-2.17 [-4.16, -0.18]
16 Mental state: 6. Specific - average score - depression (SDS, endpoint, high score = worse, short-term)	1	42	Mean Difference (IV, Fixed, 95% CI)	-24.25 [-28.01, -20.49]
17 Mental state: 8. Specific - not improved (auditory hallucinations, endpoint)	3		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
17.1 still existed after 30 days, short-term	1	216	Risk Ratio (M-H, Fixed, 95% CI)	0.24 [0.13, 0.44]
17.2 still existed or appeared frequently after disappeared, medium-term	1	64	Risk Ratio (M-H, Fixed, 95% CI)	0.32 [0.17, 0.61]
17.3 no change of auditory hallucinations, duration unclear	1	120	Risk Ratio (M-H, Fixed, 95% CI)	0.25 [0.03, 2.17]
18 Mental state: 9. Specific - average score - auditory hallucinations (SAHS, end- point, high score = worse) - Skewed data			Other data	No numeric data
18.1 short-term			Other data	No numeric data
18.2 medium-term			Other data	No numeric data
19 Mental state: 10. Specific - average score - hallucinations (BPRS [12th item], endpoint, high score = worse, short-term)	1	90	Mean Difference (IV, Fixed, 95% CI)	-0.73 [-1.28, -0.18]
20 Behaviour: Leaving the study early	15		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
20.1 short-term	10	870	Risk Ratio (M-H, Fixed, 95% Cl)	1.33 [0.33, 5.45]
20.2 medium-term	5	370	Risk Ratio (M-H, Fixed, 95% CI)	3.58 [0.60, 21.27]
21 Service outcomes: Time in hospital (days)	1	120	Mean Difference (IV, Fixed, 95% CI)	-16.0 [-19.54, -12.46]
22 Adverse effects: 1. General - aver- age score (TESS, endpoint, high score = worse, short-term)	1	90	Mean Difference (IV, Fixed, 95% CI)	-2.80 [-3.09, -2.51]
23 Adverse effects: 2a. Specific - ex- trapyramidal symptoms	5		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
23.1 short-term - overall	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.66 [0.40, 1.09]

Acupuncture for schizophrenia (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
23.2 short-term - specific - akathisia	1	75	Risk Ratio (M-H, Fixed, 95% CI)	0.58 [0.23, 1.48]
23.3 short-term - specific - tremor	1	75	Risk Ratio (M-H, Fixed, 95% CI)	0.73 [0.24, 2.18]
23.4 medium-term - overall	2	156	Risk Ratio (M-H, Fixed, 95% CI)	0.62 [0.32, 1.23]
23.5 medium-term - specific - myotonia	1	60	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.07, 15.26]
23.6 medium-term - specific - tremor	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.67 [0.12, 3.71]
23.7 medium-term - specific - akathisia	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.6 [0.16, 2.29]
24 Adverse effects: 2b. Specific - ex- trapyramidal symptoms -overall (short- term) - subgroup analysis	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.66 [0.40, 1.09]
24.1 traditional acupuncture	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.17 [0.02, 1.30]
24.2 electroacupuncture	2	142	Risk Ratio (M-H, Fixed, 95% CI)	0.79 [0.47, 1.34]
25 Adverse effects: 3. Specific - Central Nervous System	5		Risk Ratio (M-H, Fixed, 95% Cl)	Subtotals only
25.1 medium-term - anxiety	1	96	Risk Ratio (M-H, Fixed, 95% CI)	0.42 [0.09, 2.05]
25.2 short-term - insomnia	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.30 [0.11, 0.83]
25.3 medium-term - insomnia	2	156	Risk Ratio (M-H, Fixed, 95% CI)	0.34 [0.12, 1.02]
25.4 medium-term - headache	1	60	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [0.33, 27.23]
26 Adverse effects: 4. Specific - anticholin- ergic symptoms	5		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
26.1 short-term - dry mouth	3	202	Risk Ratio (M-H, Fixed, 95% CI)	1.18 [0.47, 3.00]
26.2 short-term - blurred vision	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.24, 3.24]
26.3 medium-term - blurred vision	2	156	Risk Ratio (M-H, Fixed, 95% CI)	1.0 [0.15, 6.64]

Acupuncture for schizophrenia (Review)



Cochrane Database of Systematic Reviews

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
26.4 medium-term - sweating	2	156	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [0.13, 70.83]
26.5 short-term - constipation	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.19, 4.06]
26.6 short-term - nausea & vomiting	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.41 [0.12, 1.32]
27 Adverse effects: 5. Specific - gastroin- testinal system	4		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
27.1 medium-term - unspecified gastroin- testinal symptoms	1	96	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.10, 2.71]
27.2 short-term - constipation	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.19, 4.06]
27.3 short-term - nausea & vomiting	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.41 [0.12, 1.32]
28 Adverse effects: 6. Specific - cardiovas- cular symptoms (or headache)	6		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
28.1 short-term - dizziness	3	202	Risk Ratio (M-H, Fixed, 95% CI)	0.94 [0.32, 2.75]
28.2 medium-term - dizziness or headache	1	96	Risk Ratio (M-H, Fixed, 95% CI)	0.35 [0.04, 3.22]
28.3 short-term - tachycardia	3	217	Risk Ratio (M-H, Fixed, 95% CI)	0.87 [0.42, 1.79]
28.4 medium-term - tachycardia	2	156	Risk Ratio (M-H, Fixed, 95% CI)	0.34 [0.04, 3.20]
29 Adverse effects: 7a. Specific - metabol- ic system	4		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
29.1 short-term - weight gain	3	202	Risk Ratio (M-H, Random, 95% CI)	0.94 [0.02, 37.33]
29.2 medium-term - weight gain	1	96	Risk Ratio (M-H, Random, 95% CI)	0.21 [0.01, 4.23]
30 Adverse effects: 7b. Specific - metabol- ic system - weight gain (short-term) - sub- group analysis	3		Risk Ratio (M-H, Random, 95% CI)	Subtotals only
30.1 traditional acupuncture	1	60	Risk Ratio (M-H, Random, 95% CI)	0.14 [0.01, 2.65]
30.2 electroacupuncture	2	142	Risk Ratio (M-H, Random, 95% CI)	6.15 [0.33, 115.01]

Acupuncture for schizophrenia (Review)



Cochrane Database of Systematic Reviews

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
31 Adverse effects: 8. Specific - endocrine system	2		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
31.1 short-term - irregular menstruation	2	127	Risk Ratio (M-H, Fixed, 95% Cl)	0.33 [0.01, 7.87]
32 Adverse effects: 9. Specific - lab test	5		Risk Ratio (M-H, Fixed, 95% Cl)	Subtotals only
32.1 short-term - liver function abnormal	4	292	Risk Ratio (M-H, Fixed, 95% CI)	1.01 [0.26, 3.90]
32.2 medium-term - liver function abnor- mal	1	96	Risk Ratio (M-H, Fixed, 95% CI)	0.70 [0.12, 3.98]
32.3 short-term - ECG abnormal (myocar- dial ischaemia)	4	292	Risk Ratio (M-H, Fixed, 95% CI)	0.5 [0.05, 5.32]
32.4 short-term - blood routine test ab- normal (leukocyte change)	4	292	Risk Ratio (M-H, Fixed, 95% CI)	1.33 [0.32, 5.62]

# Analysis 1.1. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 1 Global state: Not improved, endpoint.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% CI
1.1.1 medium-term (various simila	ar criteria)				
traditional - Liu 2010	16/50	31/50		45.59%	0.52[0.33,0.82]
traditional - Luo 2006	2/30	5/30		7.35%	0.4[0.08,1.9]
traditional - Tang 2005	9/42	32/42		47.06%	0.28[0.15,0.51]
Subtotal (95% CI)	122	122	◆	100%	0.4[0.28,0.57]
Total events: 27 (Acupuncture adder (Standard dose antipsychotics)	d to standard dose a	ntipsychotics), 68			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =2.51, df	f=2(P=0.28); I <sup>2</sup> =20.36	%			
Test for overall effect: Z=5.08(P<0.00	001)				
1.1.2 duration unclear					
laser - Ma 1999	0/60	2/60		100%	0.2[0.01,4.08]
Subtotal (95% CI)	60	60		100%	0.2[0.01,4.08]
Total events: 0 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 2			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.05(P=0.3)					
Test for subgroup differences: Chi <sup>2</sup> =	0.2, df=1 (P=0.66), I <sup>2</sup> =	-0%			
Favours Acupuncture	added to standard d	ose antipsychotics	0.01 0.1 1 10	<sup>100</sup> Favours Standard do	se antipsychotics

## Analysis 1.2. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Mental state: 1a. General - average score (BPRS, endpoint, high score = worse).

Study or subgroup	Acupun to sta antip	cture added ndard dose osychotics	Standard dose antipsychotics		Mean Difference	Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
1.2.1 short-term							
electro - Ding 2005	25	45 (9.1)	25	49.9 (9.4)		3.52%	-4.89[-10.02,0.24]
laser - Ma 1999	60	26.4 (3.9)	60	31.3 (4)		46.21%	-4.89[-6.31,-3.47]
traditional - Liu 2010	48	19.8 (3.5)	49	23.5 (4)		41.46%	-3.67[-5.17,-2.17]
traditional - Luo 2006	30	30.7 (6.5)	30	34.8 (6.4)		8.81%	-4.16[-7.41,-0.91]
Subtotal ***	163		164		◆	100%	-4.32[-5.28,-3.36]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.4, df	=3(P=0.7);	I <sup>2</sup> =0%					
Test for overall effect: Z=8.79(P<0.0	001)						
1.2.2 medium-term							
acupoint inj - Yang 2000	34	22.8 (7.5)	30	23.5 (9.8)	+	7.76%	-0.7[-5.02,3.62]
traditional - Liu 2010	47	17.4 (3.7)	49	22.3 (3.1)		77.43%	-4.87[-6.24,-3.5]
traditional - Luo 2006	30	26.5 (6.8)	30	37.9 (5.5)	<b>↓</b>	14.81%	-11.35[-14.48,-8.22]
Subtotal ***	111		109		◆	100%	-5.51[-6.71,-4.3]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =19.01,	df=2(P<0.	0001); I <sup>2</sup> =89.48%					
Test for overall effect: Z=8.97(P<0.0	001)						
Test for subgroup differences: Chi <sup>2</sup>	2.28, df=1	(P=0.13), I <sup>2</sup> =56.0	)5%				
Favours Acu	puncture	added to standa	rd dose a	ntipsychotics	-10 -5 0 5 10	Favours Sta chotics	ndard dose antipsy-

### Analysis 1.3. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Mental state: 1b. General average score (BPRS, endpoint, high score = worse, medium-term) - subgroup analysis.

Study or subgroup	Acupun to star antip	uncture added Standard dose tandard dose antipsychotics tipsychotics		dard dose sychotics	Mean Difference	Weight	Mean Difference			
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% Cl		Random, 95% CI			
1.3.1 traditional acupuncture										
traditional - Liu 2010	47	17.4 (3.7)	49	22.3 (3.1)		52.45%	-4.87[-6.24,-3.5]			
traditional - Luo 2006	30	26.5 (6.8)	30	37.9 (5.5)	•	47.55%	-11.35[-14.48,-8.22]			
Subtotal ***	77		79		•	100%	-7.95[-14.29,-1.61]			
Heterogeneity: Tau <sup>2</sup> =19.48; Chi <sup>2</sup> =13.86, df=1(P=0); l <sup>2</sup> =92.78%										
Test for overall effect: Z=2.46(P=0.02	L)									
1.3.2 acupoint injection										
acupoint inj - Yang 2000	34	22.8 (7.5)	30	23.5 (9.8)	+	100%	-0.7[-5.02,3.62]			
Subtotal ***	34		30		<b>♦</b>	100%	-0.7[-5.02,3.62]			
Heterogeneity: Not applicable										
Test for overall effect: Z=0.32(P=0.75	5)									
Test for subgroup differences: Chi <sup>2</sup> =	3.43, df=1	(P=0.06), I <sup>2</sup> =70.8	5%							
Favours Acu	ouncture	added to standar	d dose a	ntipsychotics	-100 -50 0 50	<sup>100</sup> Favours Star chotics	ndard dose antipsy-			

## Analysis 1.4. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Mental state: 2a. General - average score (PANSS, endpoint, high score = worse).

Study or subgroup	Acupuncture added to standard dose antipsychotics		Standard dose antipsychotics		Mean Difference	Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
1.4.1 short-term							
electro - Cheng 2009	30	67.2 (10.8)	30	68.1 (11.4)	+	10.8%	-0.93[-6.54,4.68]
electro - Ding 2005	25	74.1 (11.1)	25	86 (10.2)		9.7%	-11.96[-17.88,-6.04]
electro - Wang 2005	40	46.6 (6.8)	35	51.2 (6.4)		38.03%	-4.6[-7.59,-1.61]
traditional - Ma 2008	30	51.5 (6.4)	30	50.2 (4.8)		41.47%	1.3[-1.56,4.16]
Subtotal ***	125		120		•	100%	-2.47[-4.31,-0.63]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =18.78,	df=3(P=0);	; I <sup>2</sup> =84.03%					
Test for overall effect: Z=2.63(P=0.0	1)						
1.4.2 medium-term							
electro - Cheng 2009	30	67.2 (10.8)	30	68.1 (11.4)		22.12%	-0.93[-6.54,4.68]
electro - Wang 2005	40	46.6 (6.8)	35	51.2 (6.4)		77.88%	-4.6[-7.59,-1.61]
Subtotal ***	70		65		•	100%	-3.79[-6.43,-1.15]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.28, d	lf=1(P=0.26	5); I <sup>2</sup> =21.91%					
Test for overall effect: Z=2.81(P=0)							
Test for subgroup differences: Chi <sup>2</sup>	=0.64, df=1	(P=0.42), I <sup>2</sup> =0%					
Favours Acu	puncture a	added to standa	rd dose a	ntipsychotics	-20 -10 0 10 20	Favours Sta chotics	ndard dose antipsy-

### Analysis 1.5. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Mental state: 2b. General average score (PANSS, endpoint, high score = worse, short-term) - subgroup analysis.

Study or subgroup	Acupun to sta antip	cture added ndard dose osychotics	Stan antip	dard dose osychotics	Mean Difference	Weig	ht Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Random, 95% CI		Random, 95% Cl
1.5.1 traditional acupuncture							
traditional - Ma 2008	30	51.5 (6.4)	30	50.2 (4.8)	+	100	1.3[-1.56,4.16]
Subtotal ***	30		30		•	100	1.3[-1.56,4.16]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.89(P=0.37	')						
1.5.2 electroacupuncture							
electro - Cheng 2009	30	67.2 (10.8)	30	68.1 (11.4)	+	30.43	-0.93[-6.54,4.68]
electro - Ding 2005	25	74.1 (11.1)	25	86 (10.2)	-	29.29	-11.96[-17.88,-6.04]
electro - Wang 2005	40	46.6 (6.8)	35	51.2 (6.4)		40.28	-4.6[-7.59,-1.61]
Subtotal ***	95		90		•	100	-5.64[-10.93,-0.35]
Heterogeneity: Tau <sup>2</sup> =15.78; Chi <sup>2</sup> =7.3	9, df=2(P	=0.02); I <sup>2</sup> =72.94%	)				
Test for overall effect: Z=2.09(P=0.04	)						
Test for subgroup differences: Chi <sup>2</sup> =	5.11, df=1	(P=0.02), l <sup>2</sup> =80.4	2%			l	
Favours Acup	ouncture	added to standa	rd dose a	ntipsychotics -10	0 -50 0 50	<sup>100</sup> Favo choti	urs Standard dose antipsy- cs



# Analysis 1.6. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Mental state: 2c. General - average score (PANSS, endpoint, high score = worse, short-term) - electroacupuncture subgroup analysis.

Study or subgroup	Acupuncture added to standard dose antipsychotics		Stan antip	Standard dose antipsychotics		Mean Difference			Weight	Mean Difference	
	N	Mean(SD)	Ν	Mean(SD)		F	ixed, 95% C	l			Fixed, 95% CI
electro - Cheng 2009	30	67.2 (10.8)	30	68.1 (11.4)			+			22.12%	-0.93[-6.54,4.68]
electro - Wang 2005	40	46.6 (6.8)	35	51.2 (6.4)			H			77.88%	-4.6[-7.59,-1.61]
Total ***	70		65				•			100%	-3.79[-6.43,-1.15]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.28, o	df=1(P=0.2	6); I <sup>2</sup> =21.91%									
Test for overall effect: Z=2.81(P=0)											
Favours Acupuncture added to standard dose antipsychotics						-50	0	50	100	Favours Star chotics	ndard dose antipsy-

### Analysis 1.7. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Mental state: 2d. General - average score (PANSS, endpoint, high score = worse) - Skewed data.

Mental state: 2d. General - average score (PANSS, endpoint, high score = worse) - Skewed data

Study	Intervention	Mean	SD	N	Note
			short-term		
electro - Chen 2008	Electroacupuncture added to standard dose antipsychotics	50.73	13.32	30	Compared with the score before treatment P < 0.05
electro - Chen 2008	Standard dose antipsy- chotics	72.30	7.01	30	
electro - Yao 2006	Electroacupuncture added to standard dose antipsychotics	46.51	17.10	45	Compared with the score before treatment P < 0.01
electro - Yao 2006	Standard dose antipsy- chotics	46.45	17.23	45	Compared with the score before treatment P < 0.01
traditional - Bouhlel 2011	Traditional acupuncture added to standard dose antipsychotics	77.40	25.73	15	P = 0.501
traditional - Bouhlel 2011	Standard dose antipsy- chotics	77.81	26.77	16	
			medium-term		
electro - Chen 2008	Electroacupuncture added to standard dose antipsychotics	32.02	11.21	30	Compared with the score before treatment P < 0.01
electro - Chen 2008	Standard dose antipsy- chotics	52.10	10.32	30	Compared with the score before treatment P < 0.01

# Analysis 1.8. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 8 Mental state: 2e. General - not improved (PANSS), endpoint.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio				Weight Risk Ratio	
	n/N	n/N		M-H	, Fixed, 95	% CI		M-H, Fixed, 95% Cl	
1.8.1 reduced rate < 25%, show					I				
Favours Acupunct	ure added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard dose antipsychotics	

Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% CI
electro - Chen 2008	10/32	18/30		43.36%	0.52[0.29,0.94]
electro - Wang 2005	14/40	19/35		47.3%	0.64[0.38,1.08]
traditional - Ma 2008	5/30	4/30		9.34%	1.25[0.37,4.21]
Subtotal (95% CI)	102	95	•	100%	0.65[0.45,0.94]
Total events: 29 (Acupuncture adde (Standard dose antipsychotics)	ed to standard dose a	ntipsychotics), 41			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.65, c	lf=2(P=0.44); I <sup>2</sup> =0%				
Test for overall effect: Z=2.29(P=0.0	2)				
1.8.2 reduced rate < 30%, short-to	erm				
electro - Yao 2006	10/45	11/45	- <mark></mark>	100%	0.91[0.43,1.92]
Subtotal (95% CI)	45	45	+	100%	0.91[0.43,1.92]
Total events: 10 (Acupuncture adde (Standard dose antipsychotics)	ed to standard dose a	ntipsychotics), 11			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.25(P=0.8	;)				
1.8.3 reduced rate < 30%, follow-	up, medium-term				
electro - Yao 2006	11/45	16/45	- <mark></mark> -	100%	0.69[0.36,1.31]
Subtotal (95% CI)	45	45	•	100%	0.69[0.36,1.31]
Total events: 11 (Acupuncture adde (Standard dose antipsychotics)	ed to standard dose a	ntipsychotics), 16			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.14(P=0.2	6)				
Test for subgroup differences: Chi <sup>2</sup>	=0.63, df=1 (P=0.73), I	<sup>2</sup> =0%			
Favours Acupuncture	added to standard d	lose antipsychotics 0.01	0.1 1 10	<sup>100</sup> Favours Standard de	ose antipsychotics

# Analysis 1.9. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 9 Mental state: 3. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data.

Mental state: 3. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	Ν	Note
electro - Zhang 1993	Electroacupuncture added to standard dose antipsychotics	6.71	6.15	38	P < 0.01
electro - Zhang 1993	Standard dose antipsy- chotics	10.65	6.95	31	
traditional - Bouhlel 2011	Traditional acupuncture added to standard dose antipsychotics	42.00	28.69	15	P = 0.539
traditional - Bouhlel 2011	Standard dose antipsy- chotics	46.75	25.05	16	



### Analysis 1.10. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 10 Mental state: 4a.Specific - average score - negative symptoms (SANS, endpoint, high score = worse).

Study or subgroup	Acupun to star antip	cture added ndard dose sychotics	Stano antip	dard dose sychotics	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)	Fixed, 95% Cl		Fixed, 95% CI
1.10.1 short-term							
traditional - Luo 2006	30	56.3 (11)	30	63.9 (10.3)	+	100%	-7.66[-13.05,-2.27]
Subtotal ***	30		30		•	100%	-7.66[-13.05,-2.27]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=0	(P<0.0001	); I <sup>2</sup> =100%					
Test for overall effect: Z=2.79(P=0.01	.)						
1.10.2 medium-term							
traditional - Luo 2006	30	47.6 (11.6)	30	59.9 (8.7)	+	100%	-12.35[-17.54,-7.16]
Subtotal ***	30		30		•	100%	-12.35[-17.54,-7.16]
Heterogeneity: Not applicable							
Test for overall effect: Z=4.67(P<0.00	001)						
Test for subgroup differences: Chi <sup>2</sup> =	1.51, df=1	(P=0.22), I <sup>2</sup> =33.7	5%				
Favours Acup	ouncture a	added to standar	d dose a	ntipsychotics	-100 -50 0 50 100	Favours S chotics	tandard dose antipsy-

Analysis 1.11. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 11 Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data.

Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	Ν	Note
		traditi	onal acupuncture		
traditional - Bouhlel 2011	Traditional acupuncture added to standard dose antipsychotics	55.60	24.71	15	P = 0.406
traditional - Bouhlel 2011	Standard dose antipsy- chotics	52.81	30.88	16	
		elect	roacupuncture		
electro - Zhang 1993	Electroacupuncture added to standard dose antipsychotics	24.97	20.86	38	P < 0.005
electro - Zhang 1993	Standard dose antipsy- chotics	37.26	16.02	31	

### Analysis 1.12. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 12 Mental state: 5a. Specific - average score - depression (HAMD, endpoint, high score = worse, short-term).

Study or subgroup	Acupu to sta anti	ncture added ndard dose osychotics	Stan antij	Standard dose antipsychotics		Mean Dif	ference		Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Random	, 95% CI			Random, 95% Cl
electro - Chen 2006	33	9.9 (4.3)	34	16.8 (5.7)		+			49.95%	-6.9[-9.31,-4.49]
electro - Zhang 2001	22	12.1 (3.9)	20	22.5 (4.1)		•			50.05%	-10.41[-12.81,-8.01]
Total ***	55		54			•	I		100%	-8.66[-12.1,-5.22]
Favo	urs Acupuncture	added to standa	rd dose a	ntipsychotics	-100	-50 0	50	100	Favours Sta chotics	ndard dose antipsy-

### Acupuncture for schizophrenia (Review)

Copyright  $\ensuremath{\mathbb S}$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

Cochrane Library

Trusted evidence. Informed decisions. Better health.

Cochrane Database of Systematic Reviews

Study or subgroup	Acupu to sta anti	ncture added andard dose psychotics	Sta ant	ndard dose ipsychotics		Mean Difference		Mean Difference Weight		Weight Mean Difference
	N	Mean(SD)	Ν	Mean(SD)		Rar	ndom, 95%	6 CI		Random, 95% Cl
Heterogeneity: Tau <sup>2</sup> =4.65; Ch	i²=4.08, df=1(P=	=0.04); I <sup>2</sup> =75.48%								
Test for overall effect: Z=4.93	P<0.0001)							I		
Favou	rs Acupuncture	added to standa	rd dose	antipsychotics	-100	-50	0	50	100	Favours Standard dose antipsy- chotics

### Analysis 1.13. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 13 Mental state: 5b. Specific - not improved - depression (HAMD, reduced rate < 25%, short-term).

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95	5% CI			M-H, Fixed, 95% Cl
electro - Chen 2006	4/33	25/34	_				59.5%	0.16[0.06,0.42]
electro - Zhang 2001	3/22	16/20					40.5%	0.17[0.06,0.5]
Total (95% CI)	55	54	-	◆			100%	0.17[0.08,0.34]
Total events: 7 (Acupuncture adde (Standard dose antipsychotics)	ed to standard dose an	tipsychotics), 41						
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=	=1(P=0.96); I <sup>2</sup> =0%							
Test for overall effect: Z=4.95(P<0.	0001)							
Favours Acupunctur	re added to standard d	ose antipsychotics	0.01 0	.1 1	10	100	Favours Standard do	se antipsychotics

### Analysis 1.14. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 14 Mental state: 7b. Specific not improved - auditory hallucinations (PSYRAS-AH, reduction < 20%, short-term).

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		R	isk Ratio			Weight	Risk Ratio
	n/N	n/N		м-н,	Fixed, 95	% CI			M-H, Fixed, 95% CI
electro - Cheng 2009	7/30	26/30		-	-			100%	0.27[0.14,0.52]
Total (95% CI)	30	30		•	•			100%	0.27[0.14,0.52]
Total events: 7 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 26							
Heterogeneity: Not applicable									
Test for overall effect: Z=3.88(P=0)									
Favours Acupuncture	added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard do	se antipsychotics

### Cochrane Library

# Analysis 1.15. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 15 Mental state: 7a. Specific - average score - auditory hallucinations (PSYRAS-AH, endpoint, high score = worse, short-term).

Study or subgroup	Acupun to sta antip	ncture added ndard dose osychotics	Stan antip	dard dose osychotics		Mean Difference			Weight	Mean Difference	
	Ν	Mean(SD)	Ν	Mean(SD)		I	ixed, 95% C	I			Fixed, 95% CI
electro - Cheng 2009	30	22.6 (3.6)	30	24.8 (4.3)			+			100%	-2.17[-4.16,-0.18]
Total ***	30		30				•			100%	-2.17[-4.16,-0.18]
Heterogeneity: Not applicable											
Test for overall effect: Z=2.13(P=0.03	3)										
Favours Acup	ouncture	added to standa	rd dose a	ntipsychotics	-100	-50	0	50	100	Favours Sta chotics	ndard dose antipsy-

### Analysis 1.16. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 16 Mental state: 6. Specific - average score - depression (SDS, endpoint, high score = worse, short-term).

Study or subgroup	Acupun to sta antip	octure added ndard dose osychotics	Stan antip	Standard dose antipsychotics		Mean Difference				Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)			Fixed, 9	95% CI			Fixed, 95% CI
electro - Zhang 2001	22	34.5 (6.4)	20	58.8 (6)			+			100%	-24.25[-28.01,-20.49]
Total ***	22		20				•			100%	-24.25[-28.01,-20.49]
Heterogeneity: Not applicable											
Test for overall effect: Z=12.64(P<0	.0001)				1						
Favours Act	upuncture	added to standa	rd dose a	ntipsychotics	-100	-50	0		50 100	Favours Sta chotics	ndard dose antipsy-

## Analysis 1.17. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 17 Mental state: 8. Specific - not improved (auditory hallucinations, endpoint).

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Rati	io	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 9	5% CI		M-H, Fixed, 95% CI
1.17.1 still existed after 30 days, sh	ort-term					
acupoint cat - Wang 1997	11/108	46/108			100%	0.24[0.13,0.44]
Subtotal (95% CI)	108	108	•		100%	0.24[0.13,0.44]
Total events: 11 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose ai	ntipsychotics), 46				
Heterogeneity: Not applicable						
Test for overall effect: Z=4.66(P<0.00	01)					
1.17.2 still existed or appeared free um-term	quently after disap	peared, medi-				
acupoint inj - Yang 2000	8/34	22/30	- <mark></mark> -		100%	0.32[0.17,0.61]
Subtotal (95% CI)	34	30	<b>•</b>		100%	0.32[0.17,0.61]
Total events: 8 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 22				
Favours Acupuncture a	added to standard d	ose antipsychotics	0.01 0.1 1	10 100	Favours Standard do	se antipsychotics

### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio				Weight	Risk Ratio
	n/N	n/N		М-Н, F	ixed, 95%	CI			M-H, Fixed, 95% Cl
Heterogeneity: Not applicable									
Test for overall effect: Z=3.46(P=0)									
1.17.3 no change of auditory hallu laser - Ma 1999 Subtotal (95% CI)	cinations, duration 1/60 60	unclear 4/60 60	_					100% <b>100%</b>	0.25[0.03,2.17] 0.25[0.03.2.17]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 4							
Heterogeneity: Not applicable									
Test for overall effect: Z=1.26(P=0.21	)								
Test for subgroup differences: Chi <sup>2</sup> =0	0.43, df=1 (P=0.8), I <sup>2</sup> =	:0%							
Favours Acupuncture	added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard dos	se antipsychotics

# Analysis 1.18. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 18 Mental state: 9. Specific - average score - auditory hallucinations (SAHS, endpoint, high score = worse) - Skewed data.

Mental state: 9. Specific - average score - auditory hallucinations (SAHS, endpoint, high score = worse) - Skewed data
--

Study	Intervention	Mean	SD	Ν	Note					
short-term										
traditional - Liu 2010	Traditional acupuncture added to standard dose antipsychotics	9.26	5.37	48	P < 0.05					
traditional - Liu 2010	Standard dose antipsy- chotics	13.24	5.07	49						
			medium-term							
traditional - Liu 2010	Traditional acupuncture added to standard dose antipsychotics	5.12	3.64	47	P < 0.01					
traditional - Liu 2010	Standard dose antipsy- chotics	12.63	2.89	49						

# Analysis 1.19. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 19 Mental state: 10. Specific - average score - hallucinations (BPRS [12th item], endpoint, high score = worse, short-term).

Study or subgroup	Acupuncture added to standard dose antipsychotics		Standard dose antipsychotics		Mean Difference	Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
acupoint inj - Wang 2000	45	3.6 (1.4)	45	4.3 (1.2)	+	100%	-0.73[-1.28,-0.18]
Total ***	45		45		•	100%	-0.73[-1.28,-0.18]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df	=0(P<0.0001	.); I <sup>2</sup> =100%					
Test for overall effect: Z=2.62(P=0	0.01)						
Favours A	cupuncture	added to standa	rd dose a	intipsychotics	-10 -5 0 5 10	Favours Sta	ndard dose antipsy-

# Analysis 1.20. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 20 Behaviour: Leaving the study early.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% Cl
1.20.1 short-term					
acupoint cat - Wang 1997	0/108	0/108			Not estimable
acupoint inj - Wang 2000	0/45	0/45			Not estimable
electro - Chen 2006	0/33	0/34			Not estimable
electro - Cheng 2009	4/30	3/30		100%	1.33[0.33,5.45]
electro - Ding 2005	0/25	0/25			Not estimable
electro - Wang 2005	0/40	0/35			Not estimable
electro - Yao 2006	0/45	0/45			Not estimable
electro - Zhang 2001	0/22	0/20			Not estimable
laser - Ma 1999	0/60	0/60			Not estimable
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	438	432		100%	1.33[0.33,5.45]
Total events: 4 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.4(P=0.69)					
1.20.2 medium-term					
acupoint inj - Yang 2000	0/34	0/30			Not estimable
electro - Chen 2008	2/32	0/30		- 34.02%	4.7[0.23,94.01]
traditional - Liu 2010	3/50	1/50	——————————————————————————————————————	65.98%	3[0.32,27.87]
traditional - Luo 2006	0/30	0/30			Not estimable
traditional - Tang 2005	0/42	0/42			Not estimable
Subtotal (95% CI)	188	182		100%	3.58[0.6,21.27]
Total events: 5 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 1			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.06, df	=1(P=0.81); I <sup>2</sup> =0%				
Test for overall effect: Z=1.4(P=0.16)					
Test for subgroup differences: Chi <sup>2</sup> =0	0.72, df=1 (P=0.39), l <sup>2</sup>	<sup>2</sup> =0%			
Favours Acupuncture a	added to standard d	ose antipsychotics 0.01	0.1 1 10 1	<sup>00</sup> Favours Standard do	se antipsychotics

# Analysis 1.21. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 21 Service outcomes: Time in hospital (days).

Study or subgroup	Acupuncture added to standard dose antipsychotics		Standard dose antipsychotics			Mean Difference			Weight	Mean Difference	
	Ν	Mean(SD)	Ν	Mean(SD)			Fixed, 95%	CI			Fixed, 95% CI
laser - Ma 1999	60	46 (9.6)	60	62 (10.2)			+			100%	-16[-19.54,-12.46]
Total ***	60		60				•			100%	-16[-19.54,-12.46]
Heterogeneity: Not applicable											
Test for overall effect: Z=8.85(P<0.0	0001)				1						
Favours Acu	ipuncture	added to standa	rd dose a	ntipsychotics	-100	-50	0	50	100	Favours Star chotics	ndard dose antipsy-

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.



### Analysis 1.22. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 22 Adverse effects: 1. General - average score (TESS, endpoint, high score = worse, short-term).

Study or subgroup	Acupuncture added to standard dose antipsychotics		Standard dose antipsychotics		Mean Difference	Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
electro - Yao 2006	45	12.3 (0.6)	45	15.1 (0.8)	+	100%	-2.8[-3.09,-2.51]
Total ***	45		45		•	100%	-2.8[-3.09,-2.51]
Heterogeneity: Not applicable							
Test for overall effect: Z=18.78(P<0.	.0001)						
Favours Acupuncture added to standard dose antipsychotics				ntipsychotics	-5 -2.5 0 2.5 5	Favours Star chotics	ndard dose antipsy-

# Analysis 1.23. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 23 Adverse effects: 2a. Specific - extrapyramidal symptoms.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio	
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% Cl	
1.23.1 short-term - overall						
electro - Chen 2006	7/33	6/34		21.18%	1.2[0.45,3.2]	
electro - Wang 2005	11/40	15/35		57.33%	0.64[0.34,1.21]	
traditional - Ma 2008	1/30	6/30		21.5%	0.17[0.02,1.3]	
Subtotal (95% CI)	103	99	•	100%	0.66[0.4,1.09]	
Total events: 19 (Acupuncture adde (Standard dose antipsychotics)	ed to standard dose a	antipsychotics), 27				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =3.17, d	f=2(P=0.2); I <sup>2</sup> =36.979	%				
Test for overall effect: Z=1.62(P=0.1	1)					
1.23.2 short-term - specific - akat	hisia					
electro - Wang 2005	6/40	9/35		100%	0.58[0.23,1.48]	
Subtotal (95% CI)	40	35		100%	0.58[0.23,1.48]	
Total events: 6 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose ar	ntipsychotics), 9				
Heterogeneity: Not applicable						
Test for overall effect: Z=1.14(P=0.2	5)					
1.23.3 short-term - specific - trem	or					
electro - Wang 2005	5/40	6/35		100%	0.73[0.24,2.18]	
Subtotal (95% CI)	40	35		100%	0.73[0.24,2.18]	
Total events: 5 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose ar	ntipsychotics), 6				
Heterogeneity: Not applicable						
Test for overall effect: Z=0.56(P=0.5	7)					
1.23.4 medium-term - overall						
electro - Chen 2008	6/30	9/30	— <b>—</b> —	50.53%	0.67[0.27,1.64]	
traditional - Liu 2010	5/47	9/49		49.47%	0.58[0.21,1.6]	
Subtotal (95% CI)	77	79		100%	0.62[0.32,1.23]	
Favours Acupuncture	added to standard o	dose antipsychotics 0.01	0.1 1 10	<sup>100</sup> Favours Standard do	ose antipsychotics	

Acupuncture for schizophrenia (Review)
Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
Total events: 11 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose a	intipsychotics), 18			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.04, df	=1(P=0.84); I <sup>2</sup> =0%				
Test for overall effect: Z=1.37(P=0.17	)				
1.23.5 medium-term - specific - my	otonia	4 /00		1000/	
electro - Chen 2008	1/30	1/30		100%	1[0.07,15.26]
Subtotal (95% CI)	30	30		100%	1[0.07,15.26]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	itipsychotics), 1			
Heterogeneity: Not applicable					
Test for overall effect: Not applicable	2				
1.23.6 medium-term - specific - tre	mor				
electro - Chen 2008	2/30	3/30		100%	0.67[0.12,3.71]
Subtotal (95% CI)	30	30		100%	0.67[0.12,3.71]
Total events: 2 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.46(P=0.64	)				
1.23.7 medium-term - specific - ak	athisia				
electro - Chen 2008	3/30	5/30		100%	0.6[0.16.2.29]
Subtotal (95% CI)	30	30		100%	0.6[0.16.2.29]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 5			- / -
Heterogeneity: Not applicable					
Test for overall effect: Z=0.75(P=0.45	)				
Test for subgroup differences: Chi <sup>2</sup> =0	).22, df=1 (P=1), I <sup>2</sup> =0	%			
Favours Acupuncture a	added to standard d	lose antipsychotics 0.0	1 0.1 1 10 1	<sup>100</sup> Favours Standard do	se antipsychotics

### Analysis 1.24. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 24 Adverse effects: 2b. Specific - extrapyramidal symptoms -overall (short-term) - subgroup analysis.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		M-H, Fix	ed, 95% C	I			M-H, Fixed, 95% Cl
1.24.1 traditional acupuncture									
traditional - Ma 2008	1/30	6/30		•	+			21.5%	0.17[0.02,1.3]
Subtotal (95% CI)	30	30			+			21.5%	0.17[0.02,1.3]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 6							
Heterogeneity: Not applicable									
Test for overall effect: Z=1.71(P=0.09	)								
1.24.2 electroacupuncture									
electro - Chen 2006	7/33	6/34			•	1		21.18%	1.2[0.45,3.2]
Favours Acupuncture	added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard dos	se antipsychotics

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture Standard dose Risk Ratio added to stan- antipsychotics dard dose an- tipsychotics				Risk Ratio		Risk Ratio		Risk Ratio		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		M-I	H, Fixed, 95%	СІ			M-H, Fixed, 95% Cl							
electro - Wang 2005	11/40	15/35						57.33%	0.64[0.34,1.21]							
Subtotal (95% CI)	73	69			•			78.5%	0.79[0.47,1.34]							
Total events: 18 (Acupuncture addec (Standard dose antipsychotics)	d to standard dose ai	ntipsychotics), 21														
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.12, df	=1(P=0.29); I <sup>2</sup> =10.999	%														
Test for overall effect: Z=0.86(P=0.39	)															
Total (95% CI)	103	99			•			100%	0.66[0.4,1.09]							
Total events: 19 (Acupuncture addec (Standard dose antipsychotics)	d to standard dose ai	ntipsychotics), 27														
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =3.17, df	=2(P=0.2); I <sup>2</sup> =36.97%	1														
Test for overall effect: Z=1.62(P=0.11	)															
Test for subgroup differences: Chi <sup>2</sup> =2	2.07, df=1 (P=0.15), l <sup>2</sup>	=51.8%														
Favours Acupuncture a	added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard do	se antipsychotics							

# Analysis 1.25. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 25 Adverse effects: 3. Specific - Central Nervous System.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
1.25.1 medium-term - anxiety					
traditional - Liu 2010	2/47	5/49		100%	0.42[0.09,2.05]
Subtotal (95% CI)	47	49		100%	0.42[0.09,2.05]
Total events: 2 (Acupuncture added t (Standard dose antipsychotics)	to standard dose an	tipsychotics), 5			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.08(P=0.28)	)				
1.25.2 short-term - insomnia					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	4/40	9/35		68.09%	0.39[0.13,1.15]
traditional - Ma 2008	0/30	4/30		31.91%	0.11[0.01,1.98]
Subtotal (95% CI)	103	99	$\bullet$	100%	0.3[0.11,0.83]
Total events: 4 (Acupuncture added t (Standard dose antipsychotics)	to standard dose an	tipsychotics), 13			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.68, df	=1(P=0.41); I <sup>2</sup> =0%				
Test for overall effect: Z=2.33(P=0.02)	)				
1.25.3 medium-term - insomnia					
electro - Chen 2008	1/30	5/30	<b>_</b>	42.18%	0.2[0.02,1.61]
traditional - Liu 2010	3/47	7/49		57.82%	0.45[0.12,1.63]
Subtotal (95% CI)	77	79		100%	0.34[0.12,1.02]
Total events: 4 (Acupuncture added t (Standard dose antipsychotics)	to standard dose an	tipsychotics), 12			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.42, df	=1(P=0.52); I <sup>2</sup> =0%				
Test for overall effect: Z=1.93(P=0.05)	)				
				I	
Favours Acupuncture a	added to standard d	ose antipsychotics 0.0	1 0.1 1 10	<sup>100</sup> Favours Standard do	se antipsychotics

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio		Risk Ratio		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95%	CI			M-H, Fixed, 95% Cl				
1.25.4 medium-term - headad	che											
electro - Chen 2008	3/30	1/30				-	100%	3[0.33,27.23]				
Subtotal (95% CI)	30	30					100%	3[0.33,27.23]				
Total events: 3 (Acupuncture a (Standard dose antipsychotics	dded to standard dose an )	tipsychotics), 1										
Heterogeneity: Not applicable												
Test for overall effect: Z=0.98(P	9=0.33)											
Test for subgroup differences:	Chi <sup>2</sup> =3.59, df=1 (P=0.31), I	<sup>2</sup> =16.54%										
Favours Acupund	cture added to standard d	lose antipsychotics	0.01 0.1	1	10	100	Favours Standard do	ose antipsychotics				

# Analysis 1.26. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 26 Adverse effects: 4. Specific - anticholinergic symptoms.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% CI
1.26.1 short-term - dry mouth					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	6/40	4/35		58.72%	1.31[0.4,4.28]
traditional - Ma 2008	3/30	3/30	<b>-</b>	41.28%	1[0.22,4.56]
Subtotal (95% CI)	103	99		100%	1.18[0.47,3]
Total events: 9 (Acupuncture added (Standard dose antipsychotics)	to standard dose ar	tipsychotics), 7			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.08, df	=1(P=0.78); I <sup>2</sup> =0%				
Test for overall effect: Z=0.35(P=0.72	)				
1.26.2 short-term - blurred vision					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	4/40	4/35		100%	0.88[0.24,3.24]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	103	99		100%	0.88[0.24,3.24]
Total events: 4 (Acupuncture added (Standard dose antipsychotics)	to standard dose ar	tipsychotics), 4			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.2(P=0.84)					
1.26.3 medium-term - blurred visio	on				
electro - Chen 2008	2/30	2/30	·	100%	1[0.15,6.64]
traditional - Liu 2010	0/47	0/49			Not estimable
Subtotal (95% CI)	77	79		100%	1[0.15,6.64]
Total events: 2 (Acupuncture added (Standard dose antipsychotics)	to standard dose ar	tipsychotics), 2			
Heterogeneity: Not applicable					
Test for overall effect: Not applicable	2				
1.26.4 medium-term - sweating					
electro - Chen 2008	1/30	0/30 -		100%	3[0.13,70.83]
Favours Acupuncture a	added to standard c	lose antipsychotics 0.1	0.2 0.5 1 2 5	<sup>10</sup> Favours Standard do	ose antipsychotics

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
traditional - Liu 2010	0/47	0/49			Not estimable
Subtotal (95% CI)	77	79		100%	3[0.13,70.83]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 0			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.68(P=0.5)					
1.26.5 short-term - constipation					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	3/40	3/35		100%	0.88[0.19,4.06]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	103	99		100%	0.88[0.19,4.06]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.17(P=0.86	)				
1.26.6 short-term - nausea & vomit	ting				
electro - Chen 2006	3/33	5/34		56.92%	0.62[0.16,2.38]
electro - Wang 2005	0/40	3/35 🔶	• · · · · · · · · · · · · · · · · · · ·	43.08%	0.13[0.01,2.35]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	103	99		100%	0.41[0.12,1.32]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 8			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.99, df	=1(P=0.32); I <sup>2</sup> =0%				
Test for overall effect: Z=1.5(P=0.13)					
Test for subgroup differences: Chi <sup>2</sup> =2	2.64, df=1 (P=0.76), l	2=0%			
Favours Acupuncture a	added to standard d	ose antipsychotics 0.2	L 0.2 0.5 1 2 5 1	<sup>10</sup> Favours Standard do	ose antipsychotics

# Analysis 1.27. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 27 Adverse effects: 5. Specific - gastrointestinal system.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% Cl
1.27.1 medium-term - unspecified	gastrointestinal sy	mptoms			
traditional - Liu 2010	2/47	4/49	— <u> </u>	100%	0.52[0.1,2.71]
Subtotal (95% CI)	47	49		100%	0.52[0.1,2.71]
Total events: 2 (Acupuncture added t (Standard dose antipsychotics)	to standard dose ant	tipsychotics), 4			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.77(P=0.44)	1				
1.27.2 short-term - constipation					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	3/40	3/35		100%	0.88[0.19,4.06]
traditional - Ma 2008	0/30	0/30			Not estimable
Favours Acupuncture a	dded to standard de	ose antipsychotics	0.01 0.1 1 10	<sup>100</sup> Favours Standard dos	e antipsychotics

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95% C	I		M-H, Fixed, 95% CI
Subtotal (95% CI)	103	99				100%	0.88[0.19,4.06]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	ipsychotics), 3					
Heterogeneity: Not applicable							
Test for overall effect: Z=0.17(P=0.86)	)						
1.27.3 short-term - nausea & vomit	ting						
electro - Chen 2006	3/33	5/34				56.92%	0.62[0.16,2.38]
electro - Wang 2005	0/40	3/35				43.08%	0.13[0.01,2.35]
traditional - Ma 2008	0/30	0/30					Not estimable
Subtotal (95% CI)	103	99				100%	0.41[0.12,1.32]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	ipsychotics), 8					
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.99, df	=1(P=0.32); I <sup>2</sup> =0%						
Test for overall effect: Z=1.5(P=0.13)							
Test for subgroup differences: Chi <sup>2</sup> =0	0.61, df=1 (P=0.74), I <sup>2</sup>	=0%					
Favours Acupuncture a	added to standard do	ose antipsychotics	0.01 0.	1 1	10 100 Fa	wours Standard dos	e antipsychotics

# Analysis 1.28. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 28 Adverse effects: 6. Specific - cardiovascular symptoms (or headache).

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
1.28.1 short-term - dizziness					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	3/40	3/35		51.61%	0.88[0.19,4.06]
traditional - Ma 2008	3/30	3/30	<b>+</b>	48.39%	1[0.22,4.56]
Subtotal (95% CI)	103	99		100%	0.94[0.32,2.75]
Total events: 6 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 6			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.01, d	f=1(P=0.9); I <sup>2</sup> =0%				
Test for overall effect: Z=0.12(P=0.9)					
1.28.2 medium-term - dizziness or	rheadache				
traditional - Liu 2010	1/47	3/49		100%	0.35[0.04,3.22]
Subtotal (95% CI)	47	49		100%	0.35[0.04,3.22]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.93(P=0.35	5)				
1.28.3 short-term - tachycardia					
electro - Chen 2006	4/33	5/34		35.37%	0.82[0.24,2.8]
electro - Yao 2006	6/45	5/45		35.91%	1.2[0.39,3.65]
traditional - Ma 2008	2/30	4/30		28.72%	0.5[0.1,2.53]
Subtotal (95% CI)	108	109		100%	0.87[0.42,1.79]
Favours Acupuncture	added to standard d	ose antipsychotics	0.1 0.2 0.5 1 2 5 10	Favours Standard do	se antipsychotics

#### Acupuncture for schizophrenia (Review)

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio			Weight	Risk Ratio
	n/N	n/N	M-H	l, Fixed, 95% Cl			M-H, Fixed, 95% Cl
Total events: 12 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose ar	ntipsychotics), 14					
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.78, df	=2(P=0.68); I <sup>2</sup> =0%						
Test for overall effect: Z=0.39(P=0.7)							
1.28.4 medium-term - tachycardia							
electro - Chen 2008	0/30	1/30	•		_	50.52%	0.33[0.01,7.87]
traditional - Liu 2010	0/47	1/49			_	49.48%	0.35[0.01,8.32]
Subtotal (95% CI)	77	79				100%	0.34[0.04,3.2]
Total events: 0 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	tipsychotics), 2					
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=1(	P=0.99); I <sup>2</sup> =0%						
Test for overall effect: Z=0.94(P=0.35	)						
Test for subgroup differences: Chi <sup>2</sup> =1	L.22, df=1 (P=0.75), I <sup>2</sup>	=0%			1		
Favours Acupuncture a	added to standard do	ose antipsychotics	0.1 0.2	0.5 1 2 5	10 F	avours Standard dos	se antipsychotics

# Analysis 1.29. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 29 Adverse effects: 7a. Specific - metabolic system.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Ranc	lom, 95% Cl			M-H, Random, 95% CI
1.29.1 short-term - weight gain								
electro - Chen 2006	0/33	0/34						Not estimable
electro - Wang 2005	3/40	0/35			<b>—</b>	$\rightarrow$	49.96%	6.15[0.33,115.01]
traditional - Ma 2008	0/30	3/30	◀—		<u> </u>		50.04%	0.14[0.01,2.65]
Subtotal (95% CI)	103	99	_			-	100%	0.94[0.02,37.33]
Total events: 3 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose an	tipsychotics), 3						
Heterogeneity: Tau <sup>2</sup> =4.85; Chi <sup>2</sup> =3.18	8, df=1(P=0.07); I <sup>2</sup> =68	.52%						
Test for overall effect: Z=0.04(P=0.9	7)							
1.29.2 medium-term - weight gair	n							
traditional - Liu 2010	0/47	2/49					100%	0.21[0.01,4.23]
Subtotal (95% CI)	47	49					100%	0.21[0.01,4.23]
Total events: 0 (Acupuncture addec (Standard dose antipsychotics)	l to standard dose an	tipsychotics), 2						
Heterogeneity: Not applicable								
Test for overall effect: Z=1.02(P=0.3	1)							
Test for subgroup differences: Chi <sup>2</sup> =	=0.38, df=1 (P=0.54), l	<sup>2</sup> =0%						
Favours Acupuncture	added to standard d	lose antipsychotics	0.01	0.1	1 10	100	Favours Standard d	ose antipsychotics

#### Cochrane Library Trusted evidence. Informed decisions. Better health.

### Analysis 1.30. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 30 Adverse effects: 7b. Specific - metabolic system - weight gain (short-term) - subgroup analysis.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Random, 95% Cl		M-H, Random, 95% CI
1.30.1 traditional acupuncture					
traditional - Ma 2008	0/30	3/30	<b>↓</b>	100%	0.14[0.01,2.65]
Subtotal (95% CI)	30	30		100%	0.14[0.01,2.65]
Total events: 0 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	ipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.31(P=0.19)	)				
1.30.2 electroacupuncture					
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	3/40	0/35		100%	6.15[0.33,115.01]
Subtotal (95% CI)	73	69		- 100%	6.15[0.33,115.01]
Total events: 3 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	ipsychotics), 0			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.22(P=0.22)	)				
Test for subgroup differences: Chi <sup>2</sup> =3	8.18, df=1 (P=0.07), I <sup>2</sup>	=68.52%			
Favours Acupuncture a	added to standard do	ose antipsychotics	0.01 0.1 1 10 100	<sup>0</sup> Favours Standard do	ose antipsychotics

# Analysis 1.31. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 31 Adverse effects: 8. Specific - endocrine system.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		M-	H, Fixed, 95%	∕₀ CI			M-H, Fixed, 95% CI
1.31.1 short-term - irregular mens	struation								
electro - Chen 2006	0/33	0/34							Not estimable
traditional - Ma 2008	0/30	1/30			+			100%	0.33[0.01,7.87]
Subtotal (95% CI)	63	64						100%	0.33[0.01,7.87]
Total events: 0 (Acupuncture added (Standard dose antipsychotics)	to standard dose an	tipsychotics), 1							
Heterogeneity: Not applicable									
Test for overall effect: Z=0.68(P=0.5)									
Favours Acupuncture	added to standard d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard do	se antipsychotics

# Analysis 1.32. Comparison 1 ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 32 Adverse effects: 9. Specific - lab test.

Study or subgroup	Acupuncture added to stan- dard dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
1.32.1 short-term - liver function a	bnormal				
electro - Chen 2006	2/33	1/34		24.72%	2.06[0.2,21.65]
electro - Wang 2005	0/40	0/35			Not estimable
electro - Yao 2006	2/45	3/45	<b></b>	75.28%	0.67[0.12,3.8]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	148	144		100%	1.01[0.26,3.9]
Total events: 4 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	tipsychotics), 4			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.57, df	=1(P=0.45); I <sup>2</sup> =0%				
Test for overall effect: Z=0.02(P=0.99	)				
1.32.2 medium-term - liver functio	on abnormal				
traditional - Liu 2010	2/47	3/49		100%	0.7[0.12,3.98]
Subtotal (95% CI)	47	49		100%	0.7[0.12,3.98]
Total events: 2 (Acupuncture added (Standard dose antipsychotics)	to standard dose and	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.41(P=0.68	)				
1.32.3 short-term - ECG abnormal	(myocardial ischaer	nia)			
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	0/40	0/35			Not estimable
electro - Yao 2006	1/45	2/45	<b></b>	100%	0.5[0.05,5.32]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	148	144		100%	0.5[0.05,5.32]
Total events: 1 (Acupuncture added (Standard dose antipsychotics)	to standard dose and	tipsychotics), 2			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.57(P=0.57	)				
1.32.4 short-term - blood routine t	est abnormal (leuko	ocyte change)			
electro - Chen 2006	0/33	0/34			Not estimable
electro - Wang 2005	0/40	0/35			Not estimable
electro - Yao 2006	4/45	3/45		100%	1.33[0.32,5.62]
traditional - Ma 2008	0/30	0/30			Not estimable
Subtotal (95% CI)	148	144		100%	1.33[0.32,5.62]
Total events: 4 (Acupuncture added (Standard dose antipsychotics)	to standard dose ant	tipsychotics), 3			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.39(P=0.7)					
Test for subgroup differences: Chi <sup>2</sup> =0	0.63, df=1 (P=0.89), I <sup>2</sup>	=0%		1	
Favours Acupuncture a	added to standard de	ose antipsychotics 0.01	0.1 1 10 1	.00 Favours Standard do	se antipsychotics

### Comparison 2. ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Global state: 1. Relapse (follow-up, long- term)	1	170	Risk Ratio (M-H, Fixed, 95% CI)	0.57 [0.37, 0.89]
2 Global state: 2. Not improved (various similar criteria), endpoint (short-term)	4	272	Risk Ratio (M-H, Fixed, 95% CI)	0.83 [0.40, 1.72]
2.1 no symptoms improved obviously and insight did not recover	1	32	Risk Ratio (M-H, Fixed, 95% CI)	0.2 [0.01, 3.86]
2.2 symptoms and orientation showed no change	1	40	Risk Ratio (M-H, Fixed, 95% Cl)	0.9 [0.30, 2.68]
2.3 according to traditional criteria	2	200	Risk Ratio (M-H, Fixed, 95% Cl)	1.03 [0.35, 3.02]
3 Global state: 3a. CGI - average score - CGI- SI (endpoint, high score = worse, short- term)	1	40	Mean Difference (IV, Fixed, 95% CI)	-0.40 [-1.08, 0.28]
4 Global state: 3b. CGI - average score - CGI- GI (endpoint, high score = worse, short- term) - Skewed data			Other data	No numeric data
5 Global state: 3c. CGI - average score - CGI- EI (endpoint, high score = worse, short- term) - Skewed data			Other data	No numeric data
6 Mental state: 1a. General - average score (BPRS, endpoint, high score = worse)	5		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
6.1 short-term	5	332	Mean Difference (IV, Ran- dom, 95% CI)	-5.55 [-14.40, 3.29]
6.2 long-term	1	137	Mean Difference (IV, Ran- dom, 95% CI)	-4.87 [-8.21, -1.53]
7 Mental state: 1b. General - average score (BPRS, endpoint, high score = worse, short- term) - subgroup analysis	4	300	Mean Difference (IV, Fixed, 95% CI)	-1.36 [-3.13, 0.41]
7.1 traditional acupuncture	1	80	Mean Difference (IV, Fixed, 95% CI)	3.03 [-1.22, 7.28]
7.2 electroacupuncture	1	40	Mean Difference (IV, Fixed, 95% CI)	-4.5 [-9.35, 0.35]
7.3 acupoint injection	1	160	Mean Difference (IV, Fixed, 95% CI)	-1.87 [-4.04, 0.30]
7.4 laser acupuncture	1	20	Mean Difference (IV, Fixed, 95% CI)	-1.5 [-12.40, 9.40]
8 Mental state: 1c. General - not improved (BPRS, short-term)	3		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only

Acupuncture for schizophrenia (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
8.1 1.1 reduced rate < 30%	1	60	Risk Ratio (M-H, Fixed, 95% CI)	0.75 [0.18, 3.07]
8.2 1.2 reduced rate < 25%	1	170	Risk Ratio (M-H, Fixed, 95% Cl)	0.62 [0.31, 1.25]
8.3 1.3 reduced rate ≤ 20%	1	40	Risk Ratio (M-H, Fixed, 95% CI)	0.9 [0.30, 2.68]
9 Mental state: 1d. General - average change scores (BPRS, low score = worse, short-term)	1	60	Mean Difference (IV, Fixed, 95% CI)	0.81 [-3.17, 4.79]
10 Mental state: 2a. General - Average score (PANSS, endpoint, high score = worse, short-term) - Skew data			Other data	No numeric data
11 Mental state: 2b. General - not improved (PANSS, reduced rate < 30%, short-term)	1	80	Risk Ratio (M-H, Fixed, 95% Cl)	1.22 [0.57, 2.62]
12 Mental state: 3a. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term)	2		Mean Difference (IV, Ran- dom, 95% CI)	Subtotals only
12.1 traditional acupuncture	1	32	Mean Difference (IV, Ran- dom, 95% CI)	-13.34 [-16.94, -9.74]
12.2 acupoint catgut treatment	1	180	Mean Difference (IV, Ran- dom, 95% CI)	1.21 [0.96, 1.46]
13 Mental state: 3b. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data			Other data	No numeric data
14 Mental state: 4a. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term)	1	80	Mean Difference (IV, Fixed, 95% CI)	0.61 [-3.30, 4.52]
15 Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data			Other data	No numeric data
16 Behaviour: Leaving the study early (short-term)	8	662	Risk Ratio (M-H, Fixed, 95% CI)	0.81 [0.29, 2.29]
17 Adverse effects: 1a. General - average score (TESS, endpoint, high score = worse, short-term)	2	200	Mean Difference (IV, Fixed, 95% CI)	-0.56 [-0.86, -0.26]
18 Adverse effects: 1b. General - average score (TESS, endpoint, high score = worse, short-term) - Skewed data			Other data	No numeric data



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
19 Adverse effects: 2. Specific - average score (RESES, endpoint, high score = worse, short-term)	1	20	Mean Difference (IV, Fixed, 95% CI)	-0.6 [-0.73, -0.47]
20 Adverse effects: 3. Specific - extrapyra- midal symptoms (short-term)	3		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
20.1 overall	3	260	Risk Ratio (M-H, Fixed, 95% CI)	0.27 [0.16, 0.46]
20.2 specific - myotonia	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.21 [0.01, 4.29]
20.3 specific - tremor	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.09 [0.01, 1.69]
20.4 specific - akathisia	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.03 [0.00, 0.49]
21 Adverse effects: 4. Specific - Central Ner- vous System (short-term)	2		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
21.1 activity increasing	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.15 [0.01, 2.85]
21.2 insomnia	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.07 [0.00, 1.20]
21.3 sleepiness	2	240	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.04, 3.03]
22 Adverse effects: 5. Specific - anticholin- ergic symptoms (short-term)	2		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
22.1 dry mouth	2	240	Risk Ratio (M-H, Fixed, 95% CI)	0.22 [0.09, 0.58]
22.2 nasal congestion	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.09 [0.01, 1.69]
22.3 blurred vision	2	240	Risk Ratio (M-H, Fixed, 95% CI)	0.22 [0.06, 0.83]
22.4 constipation	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.07 [0.00, 1.20]
23 Adverse effects: 6. Specific - cardiovas- cular symptoms (short-term)	2		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
23.1 dizziness	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.70 [0.12, 4.07]
23.2 tachycardia	2	240	Risk Ratio (M-H, Fixed, 95% CI)	0.25 [0.11, 0.53]

Acupuncture for schizophrenia (Review)



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
24 Adverse effects: 7. Specific - skin infec- tion (short-term)	1	180	Risk Ratio (M-H, Fixed, 95% CI)	0.0 [0.0, 0.0]

# Analysis 2.1. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 1 Global state: 1. Relapse (follow-up, long-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95	% CI			M-H, Fixed, 95% CI
acupoint inj - Pan 2002	26/105	28/65					100%	0.57[0.37,0.89]
Total (95% CI)	105	65		•			100%	0.57[0.37,0.89]
Total events: 26 (Acupuncture addec (Standard dose antipsychotics)	I to low dose antipsy	chotics), 28						
Heterogeneity: Not applicable								
Test for overall effect: Z=2.49(P=0.01)	)							
Favours Acupunc	ture added to low d	ose antipsychotics	0.01	0.1 1	10	100	Favours Standard dos	se antipsychotics

# Analysis 2.2. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Global state: 2. Not improved (various similar criteria), endpoint (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
2.2.1 no symptoms improved obvi	ously and insight d	id not recover			
traditional - Wang 2006	0/16	2/16	+	18.64%	0.2[0.01,3.86]
Subtotal (95% CI)	16	16		18.64%	0.2[0.01,3.86]
Total events: 0 (Acupuncture added dard dose antipsychotics)	to low dose antipsy	chotics), 2 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.07(P=0.29	)				
2.2.2 symptoms and orientation sh	nowed no change				
electro - Zhou 1997	6/25	4/15	_ <b>_</b>	37.28%	0.9[0.3,2.68]
Subtotal (95% CI)	25	15	<b>•</b>	37.28%	0.9[0.3,2.68]
Total events: 6 (Acupuncture added dard dose antipsychotics)	to low dose antipsy	chotics), 4 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.19(P=0.85	)				
2.2.3 according to traditional crite	ria				
acupoint cat - Sun 2005	4/88	4/92		29.16%	1.05[0.27,4.05]
laser - Zhang 1991	2/10	2/10		14.91%	1[0.17,5.77]
Subtotal (95% CI)	98	102	• • •	44.08%	1.03[0.35,3.02]
Favours Acupund	ture added to low d	lose antipsychotics	0.002 0.1 1 10	500 Favours Standard do	se antipsychotics

Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk	Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixe	d, 95% Cl		M-H, Fixed, 95% Cl
Total events: 6 (Acupuncture added dard dose antipsychotics)	to low dose antipsy	chotics), 6 (Stan-				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=1	(P=0.97); I <sup>2</sup> =0%					
Test for overall effect: Z=0.05(P=0.96	5)					
Total (95% CI)	139	133	•		100%	0.83[0.4,1.72]
Total events: 12 (Acupuncture added (Standard dose antipsychotics)	d to low dose antips	ychotics), 12				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.07, df	f=3(P=0.79); I <sup>2</sup> =0%					
Test for overall effect: Z=0.51(P=0.61	.)					
Test for subgroup differences: Chi <sup>2</sup> =	1.05, df=1 (P=0.59),	<sup>2</sup> =0%				
			0.002 0.1	1 10 500		

Favours Acupuncture added to low dose antipsychotics 0.02 0.1 1 10 500 Favours Standard dose antipsychotics

### Analysis 2.3. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Global state: 3a. CGI - average score - CGI-SI (endpoint, high score = worse, short-term).

Study or subgroup	Acu added antij	puncture to low dose osychotics	Stan antip	dard dose osychotics		Mea	an Differe	ence		Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		Fi	xed, 95%	CI			Fixed, 95% CI
electro - Zhou 1997	25	4.6 (1)	15	5 (1.1)			+			100%	-0.4[-1.08,0.28]
Total ***	25		15				•			100%	-0.4[-1.08,0.28]
Heterogeneity: Not applicable											
Test for overall effect: Z=1.15(P=0.25	)										
Favours	Acupun	cture added to lo	w dose a	ntipsychotics	-10	-5	0	5	10	Favours Star chotics	ndard dose antipsy-

#### Analysis 2.4. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Global state: 3b. CGI average score - CGI-GI (endpoint, high score = worse, short-term) - Skewed data.

#### Global state: 3b. CGI - average score - CGI-GI (endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	N	Note
electro - Zhou 1997	Electroacupuncture added to low dose an- tipsychotics	2.2	1.1	25	No significant difference
electro - Zhou 1997	Standard dose antipsy- chotics	2.3	1.3	15	

### Analysis 2.5. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Global state: 3c. CGI average score - CGI-EI (endpoint, high score = worse, short-term) - Skewed data.

Global state: 3c. CGI - average score - CGI-EI (endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	Ν	Note
electro - Zhou 1997	Electroacupuncture added to low dose an- tipsychotics	2.5	1.3	25	P < 0.05
electro - Zhou 1997	Standard dose antipsy- chotics	1.5	0.8	15	

# Analysis 2.6. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Mental state: 1a. General - average score (BPRS, endpoint, high score = worse).

Study or subgroup	Acu added antip	puncture to low dose osychotics	Stan antip	dard dose osychotics	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
2.6.1 short-term							
acupoint inj - Pan 2002	100	28.9 (7.1)	60	30.8 (6.6)		21.54%	-1.87[-4.04,0.3]
electro - Zhou 1997	25	26.1 (6.2)	15	30.6 (8.3)		20.47%	-4.5[-9.35,0.35]
laser - Zhang 1991	10	29.3 (10.5)	10	30.8 (14.1)	+	16.39%	-1.5[-12.4,9.4]
traditional - Wang 2006	16	25.4 (4.1)	16	47.5 (7.3)		20.83%	-22.14[-26.25,-18.03]
traditional - Xu 2004	40	31.4 (9.8)	40	28.3 (9.6)	+	20.77%	3.03[-1.22,7.28]
Subtotal ***	191		141			100%	-5.55[-14.4,3.29]
Heterogeneity: Tau <sup>2</sup> =93.35; Chi <sup>2</sup> =88.	63, df=4(F	P<0.0001); I²=95.√	49%				
Test for overall effect: Z=1.23(P=0.22	2)						
2.6.2 long-term							
acupoint inj - Pan 2002	88	30.4 (10)	49	35.3 (9.3)	<b></b>	100%	-4.87[-8.21,-1.53]
Subtotal ***	88		49		◆	100%	-4.87[-8.21,-1.53]
Heterogeneity: Not applicable							
Test for overall effect: Z=2.86(P=0)							
Test for subgroup differences: Chi <sup>2</sup> =	0.02, df=1	(P=0.89), I <sup>2</sup> =0%					
Favour	s Acupuno	ture added to lo	w dose a	ntipsychotics	-20 -10 0 10 20	Favours Sta chotics	ndard dose antipsy-

# Analysis 2.7. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Mental state: 1b. General - average score (BPRS, endpoint, high score = worse, short-term) - subgroup analysis.

Study or subgroup	Acu added antip	ouncture to low dose sychotics	Stano antip	dard dose sychotics	м	ean Diffe	ence		Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		Fixed, 959	% CI			Fixed, 95% CI
2.7.1 traditional acupuncture										
traditional - Xu 2004	40	31.4 (9.8)	40	28.3 (9.6)		+•	_		17.33%	3.03[-1.22,7.28]
Subtotal ***	40		40						17.33%	3.03[-1.22,7.28]
Heterogeneity: Not applicable										
Test for overall effect: Z=1.4(P=0.16)										
2.7.2 electroacupuncture										
Favours	Acupund	ture added to lov	w dose a	ntipsychotics	-20 -10	0	10	20	Favours Star chotics	ndard dose antipsy-

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acu added antip	puncture to low dose osychotics	Stan antip	dard dose osychotics	Mean Difference	Weight	Mean Difference
	N	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
electro - Zhou 1997	25	26.1 (6.2)	15	30.6 (8.3)	-+	13.32%	-4.5[-9.35,0.35]
Subtotal ***	25		15		-	13.32%	-4.5[-9.35,0.35]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.82(P=0.07)							
2.7.3 acupoint injection							
acupoint inj - Pan 2002	100	28.9 (7.1)	60	30.8 (6.6)		66.71%	-1.87[-4.04,0.3]
Subtotal ***	100		60		•	66.71%	-1.87[-4.04,0.3]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.69(P=0.09)							
2.7.4 laser acupuncture							
laser - Zhang 1991	10	29.3 (10.5)	10	30.8 (14.1)		2.64%	-1.5[-12.4,9.4]
Subtotal ***	10		10			2.64%	-1.5[-12.4,9.4]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.27(P=0.79)							
Total ***	175		125		•	100%	-1.36[-3.13,0.41]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =5.91, df=	3(P=0.12	2); I <sup>2</sup> =49.27%					
Test for overall effect: Z=1.51(P=0.13)							
Test for subgroup differences: Chi <sup>2</sup> =5	.91, df=1	(P=0.12), I <sup>2</sup> =49.2	27%			1	
Favours	Acupund	cture added to lo	w dose a	ntipsychotics	-20 -10 0 10 2	20 Favours Sta chotics	ndard dose antipsy-

# Analysis 2.8. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 8 Mental state: 1c. General - not improved (BPRS, short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio		Weight	Risk Ratio
	n/N	n/N	M	H, Fixed, 95% Cl			M-H, Fixed, 95% CI
2.8.1 1.1 reduced rate < 30%							
electro - Cui 2000	3/30	4/30	-			100%	0.75[0.18,3.07]
Subtotal (95% CI)	30	30				100%	0.75[0.18,3.07]
Total events: 3 (Acupuncture added to dard dose antipsychotics)	o low dose antipsy	chotics), 4 (Stan-					
Heterogeneity: Not applicable							
Test for overall effect: Z=0.4(P=0.69)							
2.8.2 1.2 reduced rate < 25%							
acupoint inj - Pan 2002	13/105	13/65		- <b></b> -		100%	0.62[0.31,1.25]
Subtotal (95% CI)	105	65		•		100%	0.62[0.31,1.25]
Total events: 13 (Acupuncture added (Standard dose antipsychotics)	to low dose antipsy	ychotics), 13					
Heterogeneity: Not applicable							
Test for overall effect: Z=1.34(P=0.18)							
2.8.3 1.3 reduced rate ≤ 20%							
electro - Zhou 1997	6/25	4/15		<mark></mark>	1	100%	0.9[0.3,2.68]
Favours Acupunct	ure added to low d	ose antipsychotics	0.01 0.1	1 10	100	Favours Standard do	ose antipsychotics

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H	l, Fixed, 95%	CI			M-H, Fixed, 95% CI
Subtotal (95% CI)	25	15						100%	0.9[0.3,2.68]
Total events: 6 (Acupuncture adde dard dose antipsychotics)	d to low dose antipsyc	hotics), 4 (Stan-							
Heterogeneity: Not applicable									
Test for overall effect: Z=0.19(P=0.8	35)								
Test for subgroup differences: Chi <sup>2</sup>	=0.33, df=1 (P=0.85), I <sup>2</sup>	=0%				1	1		
Favours Acupu	ncture added to low d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard do	se antipsychotics

Favours Acupuncture added to low dose antipsychotics

#### Analysis 2.9. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 9 Mental state: 1d. General - average change scores (BPRS, low score = worse, short-term).

Study or subgroup	Acupuncture Standard dose added to low dose antipsychotics		dard dose osychotics		Mean Difference				Weight	Mean Difference	
	Ν	Mean(SD)	Ν	Mean(SD)			Fixed, 95% C	:1			Fixed, 95% CI
electro - Cui 2000	30	25 (8.5)	30	24.2 (7.2)			+			100%	0.81[-3.17,4.79]
Total ***	30		30				•			100%	0.81[-3.17,4.79]
Heterogeneity: Not applicable											
Test for overall effect: Z=0.4(P=0.69)											
Favours	Acupun	cture added to lo	w dose a	ntipsychotics	-100	-50	0	50	100	Favours Star chotics	ndard dose antipsy-

#### Analysis 2.10. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 10 Mental state: 2a. General - Average score (PANSS, endpoint, high score = worse, short-term) - Skew data.

Mental state: 2a. General - Average score (PANSS, endpoint, high score = worse, short-term) - Skew data

Study	Intervention	Mean	SD	Ν	Note
acupoint cat - Sun 2005	Acupoint catgut treat- ment added to low dose antipsychotics	52.85	15.86	88	P > 0.05
acupoint cat - Sun 2005	Standard dose antipsy- chotics	49.75	12.77	92	
electro - Xiong 2010	Electroacupuncture added to low dose an- tipsychotics	46.34	11.10	40	Cmpared with score be- fore treatment P < 0.01
electro - Xiong 2010	Standard dose antipsy- chotics	45.21	11.36	40	Cmpared with score be- fore treatment P < 0.01

#### Analysis 2.11. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 11 Mental state: 2b. General - not improved (PANSS, reduced rate < 30%, short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H, Fixed, 95	% CI			M-H, Fixed, 95% CI
electro - Xiong 2010	11/40	9/40					100%	1.22[0.57,2.62]
Total (95% CI)	40	40		+			100%	1.22[0.57,2.62]
Total events: 11 (Acupuncture added dard dose antipsychotics)	d to low dose antipsy	chotics), 9 (Stan-						
Heterogeneity: Not applicable								
Test for overall effect: Z=0.51(P=0.61	.)				1			
Favours Acupuno	cture added to low d	ose antipsychotics	0.01	0.1 1	10	100	Favours Standard dos	e antipsychotics

# Analysis 2.12. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 12 Mental state: 3a. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term).

Study or subgroup	Acu added antij	puncture to low dose osychotics	Stan antip	dard dose osychotics		Mean Difference	e	Weight	Mean Difference
	Ν	Mean(SD)	N	Mean(SD)		Random, 95% C	1		Random, 95% CI
2.12.1 traditional acupuncture									
traditional - Wang 2006	16	9 (3.4)	16	22.4 (6.5)		+		100%	-13.34[-16.94,-9.74]
Subtotal ***	16		16			♦		100%	-13.34[-16.94,-9.74]
Heterogeneity: Not applicable									
Test for overall effect: Z=7.26(P<0.00	01)								
2.12.2 acupoint catgut treatment									
acupoint cat - Sun 2005	88	9.9 (0.9)	92	8.6 (0.9)				100%	1.21[0.96,1.46]
Subtotal ***	88		92					100%	1.21[0.96,1.46]
Heterogeneity: Not applicable									
Test for overall effect: Z=9.43(P<0.00	01)								
Test for subgroup differences: Chi <sup>2</sup> =	62.42, df=	=1 (P<0.0001), I <sup>2</sup> =9	98.4%						
Favours	s Acupun	cture added to lo	w dose a	ntipsychotics	-100 -50	0	50 100	Favours Star chotics	ndard dose antipsy-

#### Analysis 2.13. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 13 Mental state: 3b. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data.

#### Mental state: 3b. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	N	Note
traditional - Xu 2004	Traditional acupuncture added to low dose an- tipsychotics	11.86	11.41	40	No significant difference
traditional - Xu 2004	Standard dose antipsy- chotics	11.12	9.87	40	



# Analysis 2.14. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 14 Mental state: 4a. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term).

Study or subgroup	Acupuncture added to low dose antipsychotics		Stan antip	dard dose osychotics	Mean Difference	Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Fixed, 95% CI		Fixed, 95% CI
traditional - Xu 2004	40	20.8 (8.4)	40	20.2 (9.5)	+	100%	0.61[-3.3,4.52]
					$\top$		
Total ***	40		40		<b>•</b>	100%	0.61[-3.3,4.52]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.31(P=0.76	)						
Favours	Acupun	cture added to lo	w dose a	ntipsychotics	-100 -50 0 50	<sup>100</sup> Favours Star chotics	idard dose antipsy-

#### Analysis 2.15. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 15 Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data.

Mental state: 4b. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	Ν	Note
traditional - Wang 2006	Traditional acupuncture added to low dose an- tipsychotics	14.23	8.68	16	P < 0.01
traditional - Wang 2006	Standard dose antipsy- chotics	30.56	13.31	16	

# Analysis 2.16. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 16 Behaviour: Leaving the study early (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
acupoint cat - Sun 2005	0/88	0/92			Not estimable
acupoint inj - Pan 2002	5/105	5/65		86.07%	0.62[0.19,2.06]
electro - Cui 2000	0/30	0/30			Not estimable
electro - Xiong 2010	0/40	0/40			Not estimable
electro - Zhou 1997	0/25	0/15			Not estimable
laser - Zhang 1991	0/10	0/10			Not estimable
traditional - Wang 2006	0/16	0/16			Not estimable
traditional - Xu 2004	2/40	1/40	+	13.93%	2[0.19,21.18]
Total (95% CI)	354	308		100%	0.81[0.29,2.29]
Total events: 7 (Acupuncture added dard dose antipsychotics)	to low dose antipsy	chotics), 6 (Stan-			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.76, df	f=1(P=0.38); I <sup>2</sup> =0%				
Test for overall effect: Z=0.39(P=0.69	))				
Favours Acupund	cture added to low d	ose antipsychotics	0.01 0.1 1 10	<sup>100</sup> Favours Standard do	se antipsychotics



#### Analysis 2.17. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 17 Adverse effects: 1a. General - average score (TESS, endpoint, high score = worse, short-term).

Study or subgroup	Acu added antip	puncture Standard dose to low dose antipsychotics ssychotics		dard dose osychotics	Mean Difference					Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		Fixed	, 95% CI				Fixed, 95% CI
acupoint inj - Pan 2002	100	3.8 (1.4)	60	4.5 (1.9)			•			31.12%	-0.7[-1.24,-0.16]
electro - Zhou 1997	25	1.4 (0.5)	15	1.9 (0.6)						68.88%	-0.5[-0.86,-0.14]
Total ***	125		75							100%	-0.56[-0.86,-0.26]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.37, df	=1(P=0.5	5); I <sup>2</sup> =0%									
Test for overall effect: Z=3.67(P=0)					1						
Favours	s Acupuno	ture added to lo	w dose a	ntipsychotics	-100	-50	0	50	100	Favours Sta chotics	ndard dose antipsy-

# Analysis 2.18. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 18 Adverse effects: 1b. General - average score (TESS, endpoint, high score = worse, short-term) - Skewed data.

Adverse effects: 1b. General - average score (TESS, endpoint, high score = worse, short-term) - Skewed data

Study	Intervention	Mean	SD	Ν	Note
electro - Xiong 2010	Electroacupuncture added to low dose an- tipsychotics	5.08	4.56	40	P < 0.01
electro - Xiong 2010	Standard dose antipsy- chotics	10.38	6.52	40	
traditional - Wang 2006	Traditional acupuncture added to low dose an- tipsychotics	2.57	3.28	16	P < 0.01
traditional - Wang 2006	Standard dose antipsy- chotics	6.93	5.28	16	
traditional - Xu 2004	Traditional acupuncture added to low dose an- tipsychotics	2.65	3.47	40	P < 0.01
traditional - Xu 2004	Standard dose antipsy- chotics	10.12	8.12	40	

### Analysis 2.19. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 19 Adverse effects: 2. Specific - average score (RESES, endpoint, high score = worse, short-term).

Study or subgroup	Acu added antip	puncture to low dose osychotics	Standard dose antipsychotics			Mean Difference				Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)		F	ixed, 95% CI				Fixed, 95% CI
laser - Zhang 1991	10	1 (0.2)	10	1.6 (0.2)						100%	-0.6[-0.73,-0.47]
Total ***	10		10							100%	-0.6[-0.73,-0.47]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=	0(P<0.0001	.); I <sup>2</sup> =100%									
Test for overall effect: Z=8.94(P<0.	0001)										
Favou	irs Acupuno	cture added to lo	w dose a	ntipsychotics	-100	-50	0	50	100	Favours Sta chotics	ndard dose antipsy-

# Analysis 2.20. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 20 Adverse effects: 3. Specific - extrapyramidal symptoms (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% CI		M-H, Fixed, 95% Cl
2.20.1 overall					
acupoint cat - Sun 2005	0/88	24/92	<b>↓</b>	52.13%	0.02[0,0.35]
electro - Cui 2000	5/30	14/30	<b>_</b>	30.46%	0.36[0.15,0.87]
laser - Zhang 1991	7/10	8/10		17.41%	0.88[0.53,1.46]
Subtotal (95% CI)	128	132	•	100%	0.27[0.16,0.46]
Total events: 12 (Acupuncture added (Standard dose antipsychotics)	to low dose antipsy	chotics), 46			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =23.67, df	=2(P<0.0001); I <sup>2</sup> =91	.55%			
Test for overall effect: Z=4.77(P<0.000	)1)				
2.20.2 specific - myotonia					
acupoint cat - Sun 2005	0/88	2/92		100%	0.21[0.01,4.29]
Subtotal (95% CI)	88	92		100%	0.21[0.01,4.29]
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 2 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.02(P=0.31)					
2.20.3 specific - tremor					
acupoint cat - Sun 2005	0/88	5/92		100%	0.09[0.01,1.69]
Subtotal (95% CI)	88	92		100%	0.09[0.01,1.69]
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 5 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.6(P=0.11)					
2.20.4 specific - akathisia					
acupoint cat - Sun 2005	0/88	17/92	<b>◀</b> <mark>···</mark>	100%	0.03[0,0.49]
Subtotal (95% CI)	88	92		100%	0.03[0,0.49]
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 17 (Stan-			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=0(F	P<0.0001); I²=100%				
Test for overall effect: Z=2.46(P=0.01)					
Test for subgroup differences: Chi <sup>2</sup> =2	.75, df=1 (P=0.43), I <sup>2</sup>	=0%			
Favours Acupunct	ure added to low d	ose antipsychotics	0.01 0.1 1 10	<sup>100</sup> Favours Standard do	ose antipsychotics

# Analysis 2.21. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 21 Adverse effects: 4. Specific - Central Nervous System (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio			Weight	Risk Ratio		
	n/N	n/N		М-Н,	Fixed, 95	% CI			M-H, Fixed, 95% Cl
2.21.1 activity increasing									
acupoint cat - Sun 2005	0/88	3/92	-					100%	0.15[0.01,2.85]
Subtotal (95% CI)	88	92				1	1	100%	0.15[0.01,2.85]
Favours Acupuncture added to low dose antipsychotics				0.1	1	10	100	Favours Standard do	se antipsychotics

#### Acupuncture for schizophrenia (Review)

Cochrane Library

Trusted evidence. Informed decisions. Better health.

Cochrane Database of Systematic Reviews

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% CI
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 3 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.26(P=0.21)					
2.21.2 insomnia					
acupoint cat - Sun 2005	0/88	7/92		100%	0.07[0,1.2]
Subtotal (95% CI)	88	92		100%	0.07[0,1.2]
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 7 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.83(P=0.07)					
2.21.3 sleepiness					
acupoint cat - Sun 2005	0/88	0/92			Not estimable
electro - Cui 2000	1/30	3/30		100%	0.33[0.04,3.03]
Subtotal (95% CI)	118	122		100%	0.33[0.04,3.03]
Total events: 1 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 3 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.98(P=0.33)					
Test for subgroup differences: Chi <sup>2</sup> =0.	74, df=1 (P=0.69), I <sup>2</sup>	=0%			
Favours Acupunct	ure added to low do	ose antipsychotics	0.01 0.1 1 10 100	Favours Standard do	se antipsychotics

# Analysis 2.22. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 22 Adverse effects: 5. Specific - anticholinergic symptoms (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk	Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixe	d, 95% CI		M-H, Fixed, 95% CI
2.22.1 dry mouth						
acupoint cat - Sun 2005	0/88	8/92	< ■		40.93%	0.06[0,1.05]
electro - Cui 2000	4/30	12/30			59.07%	0.33[0.12,0.92]
Subtotal (95% CI)	118	122	•		100%	0.22[0.09,0.58]
Total events: 4 (Acupuncture added t dard dose antipsychotics)	o low dose antipsyc	hotics), 20 (Stan-				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.41, df=	1(P=0.24); I <sup>2</sup> =28.879	%				
Test for overall effect: Z=3.09(P=0)						
2.22.2 nasal congestion						
acupoint cat - Sun 2005	0/88	5/92			100%	0.09[0.01,1.69]
Subtotal (95% CI)	88	92			100%	0.09[0.01,1.69]
Total events: 0 (Acupuncture added t dard dose antipsychotics)	o low dose antipsyc	hotics), 5 (Stan-				
Heterogeneity: Not applicable						
Test for overall effect: Z=1.6(P=0.11)						
2.22.3 blurred vision						
Favours Acupunct	ture added to low do	ose antipsychotics	0.02 0.1	1 10	50 Favours Standard o	lose antipsychotics

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% Cl
acupoint cat - Sun 2005	0/88	5/92	▲ ■	47.27%	0.09[0.01,1.69]
electro - Cui 2000	2/30	6/30	— <u>—</u> ——————————————————————————————————	52.73%	0.33[0.07,1.52]
Subtotal (95% CI)	118	122		100%	0.22[0.06,0.83]
Total events: 2 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 11 (Stan-			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.61, df=	1(P=0.43); I <sup>2</sup> =0%				
Test for overall effect: Z=2.23(P=0.03)					
2.22.4 constipation					
acupoint cat - Sun 2005	0/88	7/92		100%	0.07[0,1.2]
Subtotal (95% CI)	88	92		100%	0.07[0,1.2]
Total events: 0 (Acupuncture added to dard dose antipsychotics)	o low dose antipsyc	hotics), 7 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=1.83(P=0.07)					
Test for subgroup differences: Chi <sup>2</sup> =0.	.85, df=1 (P=0.84), I <sup>2</sup>	=0%			
Favours Acupunct	ure added to low do	ose antipsychotics	0.02 0.1 1 10 50	Favours Standard do	se antipsychotics

# Analysis 2.23. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 23 Adverse effects: 6. Specific - cardiovascular symptoms (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics	Risk Ratio	Wei	ght Risk Ratio
	n/N	n/N	M-H, Fixed, 95%	CI	M-H, Fixed, 95% Cl
2.23.1 dizziness					
acupoint cat - Sun 2005	2/88	3/92			100% 0.7[0.12,4.07]
Subtotal (95% CI)	88	92			100% 0.7[0.12,4.07]
Total events: 2 (Acupuncture added t dard dose antipsychotics)	o low dose antipsy	chotics), 3 (Stan-			
Heterogeneity: Not applicable					
Test for overall effect: Z=0.4(P=0.69)					
2.23.2 tachycardia					
acupoint cat - Sun 2005	2/88	16/92			54.62% 0.13[0.03,0.55]
electro - Cui 2000	5/30	13/30			45.38% 0.38[0.16,0.94]
Subtotal (95% CI)	118	122	<b>•</b>		100% 0.25[0.11,0.53]
Total events: 7 (Acupuncture added t dard dose antipsychotics)	o low dose antipsy	chotics), 29 (Stan-			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =1.69, df=	1(P=0.19); I <sup>2</sup> =40.87	%			
Test for overall effect: Z=3.58(P=0)					
Test for subgroup differences: Chi <sup>2</sup> =1	.13, df=1 (P=0.29), l <sup>2</sup>	2=11.12%			
Favours Acupunct	ture added to low d	ose antipsychotics	0.01 0.1 1	10 100 Favours	Standard dose antipsychotics

# Analysis 2.24. Comparison 2 ACUPUNCTURE added to LOW DOSE ANTIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 24 Adverse effects: 7. Specific - skin infection (short-term).

Study or subgroup	Acupuncture added to low dose an- tipsychotics	Standard dose antipsychotics		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		M-H	l, Fixed, 95	% CI			M-H, Fixed, 95% CI
acupoint cat - Sun 2005	0/88	0/92							Not estimable
Total (95% CI)	88	92							Not estimable
Total events: 0 (Acupuncture added t dard dose antipsychotics)	to low dose antipsyc	hotics), 0 (Stan-							
Heterogeneity: Not applicable									
Test for overall effect: Not applicable	•								
Favours Acupunc	ture added to low d	ose antipsychotics	0.01	0.1	1	10	100	Favours Standard do	se antipsychotics

#### Comparison 3. ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Global state: Not improved, endpoint (short- term)	5		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
1.1 various similar criteria	3	141	Risk Ratio (M-H, Fixed, 95% CI)	0.88 [0.53, 1.48]
1.2 talk, behaviour and expression still existed as before + reduction of BPRS < 29%	2	240	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.19, 0.59]
2 Mental state: 1. General - average score (BPRS, endpoint, high score = worse, short- term)	2		Mean Difference (IV, Random, 95% CI)	Subtotals only
2.1 traditional acupuncture	1	32	Mean Difference (IV, Random, 95% CI)	-11.56 [-16.36, -6.76]
2.2 laser acupuncture	1	21	Mean Difference (IV, Random, 95% CI)	-1.07 [-11.19, 9.05]
3 Mental state: 2. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term)	1	32	Mean Difference (IV, Fixed, 95% CI)	-5.24 [-9.06, -1.42]
4 Mental state: 3. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term)	1	32	Mean Difference (IV, Fixed, 95% CI)	-7.92 [-17.01, 1.17]
5 Behaviour: Leaving the study early (short- term)	6	421	Risk Ratio (M-H, Fixed, 95% CI)	0.0 [0.0, 0.0]
6 Adverse effects: 1. General - average scores (TESS, endpoint, short-term) - Skewed data			Other data	No numeric data



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
7 Adverse effects: 2. Specific - extrapyramidal symptoms (short-term)	1	21	Risk Ratio (M-H, Fixed, 95% CI)	0.05 [0.00, 0.83]
8 Adverse effects: 3. Specific - numbness over the ear, upper extremity and chest on the treated side (short-term)	1	40	Risk Ratio (M-H, Fixed, 95% CI)	3.0 [0.13, 69.52]

#### Analysis 3.1. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 1 Global state: Not improved, endpoint (short-term).

Study or subgroup	Acupuncture	Standard dose antipsychotics	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 95% Cl		M-H, Fixed, 95% Cl
3.1.1 various similar criteria					
electro - Zhang 1987	16/43	17/45		80.22%	0.98[0.57,1.69]
laser - Zhang 1991	1/11	2/10	+	10.12%	0.45[0.05,4.28]
traditional - Wang 2006	1/16	2/16		9.66%	0.5[0.05,4.98]
Subtotal (95% CI)	70	71	<b>•</b>	100%	0.88[0.53,1.48]
Total events: 18 (Acupuncture), 21 (S	tandard dose antip	sychotics)			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.73, df=	2(P=0.69); I <sup>2</sup> =0%				
Test for overall effect: Z=0.47(P=0.64)					
3.1.2 talk, behaviour and expressio of BPRS < 29%	n still existed as b	efore + reduction			
traditional - Zhao 2005a	10/90	10/30	— <b>—</b> —	52.63%	0.33[0.15,0.72]
traditional - Zhao 2005b	9/90	9/30	— <b>—</b> —	47.37%	0.33[0.15,0.76]
Subtotal (95% CI)	180	60	•	100%	0.33[0.19,0.59]
Total events: 19 (Acupuncture), 19 (S	tandard dose antip	sychotics)			
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0, df=1(F	P=1); I <sup>2</sup> =0%				
Test for overall effect: Z=3.81(P=0)					
Test for subgroup differences: Chi <sup>2</sup> =6	.29, df=1 (P=0.01), l	2=84.09%			
	Fa	vours Acupuncture 0.0	01 0.1 1 10 1	<sup>.00</sup> Favours Standard do	se antipsychotics

# Analysis 3.2. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 2 Mental state: 1. General - average score (BPRS, endpoint, high score = worse, short-term).

Study or subgroup	Acu	puncture	Standard dose antipsychotics		Mean Difference		Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)	Rano	lom, 95% CI		Random, 95% Cl
3.2.1 traditional acupuncture								
traditional - Wang 2006	16	36 (6.5)	16	47.5 (7.3)	<b>↓</b>		100%	-11.56[-16.36,-6.76]
Subtotal ***	16		16				100%	-11.56[-16.36,-6.76]
Heterogeneity: Not applicable								
Test for overall effect: Z=4.72(P<0.000	1)							
3.2.2 laser acupuncture								
laser - Zhang 1991	11	29.7 (8.6)	10	30.8 (14.1)			- 100%	-1.07[-11.19,9.05]
			Favours	Acupuncture	-10 -5	0 5	10 Favours s chotics	Standard dose antipsy-

#### Acupuncture for schizophrenia (Review)



Study or subgroup	Αсι	puncture	icture Standard dose antipsychotics		Mean Difference				Weight	Mean Difference	
	Ν	Mean(SD)	Ν	Mean(SD)		Rand	lom, 9	5% CI			Random, 95% CI
Subtotal ***	11		10							100%	-1.07[-11.19,9.05]
Heterogeneity: Not applicable											
Test for overall effect: Z=0.21(P=0.84)											
Test for subgroup differences: Chi <sup>2</sup> =3	.37, df=	1 (P=0.07), I <sup>2</sup> =70.32	2%								
			Favours	Acupuncture	-10	-5	0	5	10	Favours Star chotics	dard dose antipsy-

# Analysis 3.3. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 3 Mental state: 2. Specific - average score - positive symptoms (SAPS, endpoint, high score = worse, short-term).

Study or subgroup	Acup	ouncture	Standard dose antipsychotics		Mean Difference				Weight	Mean Difference	
	Ν	Mean(SD)	Ν	Mean(SD)		Fixed	, 95%	CI			Fixed, 95% CI
traditional - Wang 2006	16	17.1 (4.3)	16	22.4 (6.5)		+				100%	-5.24[-9.06,-1.42]
Total ***	16		16							100%	-5.24[-9.06,-1.42]
Heterogeneity: Not applicable											
Test for overall effect: Z=2.69(P=0.01)						1					
			Favours	Acupuncture	-10	-5	0	5	10	Favours Stan chotics	dard dose antipsy-

# Analysis 3.4. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 4 Mental state: 3. Specific - average score - negative symptoms (SANS, endpoint, high score = worse, short-term).

Study or subgroup	Acuj	ouncture	Stano antip	dard dose sychotics	Mean Difference			nce		Weight	Mean Difference
	Ν	Mean(SD)	Ν	Mean(SD)			Fixed, 95%	CI			Fixed, 95% CI
traditional - Wang 2006	16	22.6 (12.9)	16	30.6 (13.3)			+			100%	-7.92[-17.01,1.17]
Total ***	16		16				•			100%	-7.92[-17.01,1.17]
Heterogeneity: Not applicable											
Test for overall effect: Z=1.71(P=0.09)											
			Favours	Acupuncture	-100	-50	0	50	100	Favours Star chotics	ndard dose antipsy-

### Analysis 3.5. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 5 Behaviour: Leaving the study early (short-term).

Study or subgroup	Acupuncture	Standard dose antipsychotics	F	isk Ratio		Weight	Risk Ratio
	n/N	n/N	М-Н,	Fixed, 95% CI			M-H, Fixed, 95% Cl
electro - Zhang 1987	0/43	0/45					Not estimable
laser - Liu 1986	0/20	0/20					Not estimable
laser - Zhang 1991	0/11	0/10					Not estimable
traditional - Wang 2006	0/16	0/16					Not estimable
traditional - Zhao 2005a	0/90	0/30					Not estimable
traditional - Zhao 2005b	0/90	0/30			1		Not estimable
	Fav	ours Acupuncture	0.01 0.1	1 10	100	Favours Standard dos	e antipsychotics

Acupuncture for schizophrenia (Review)



Study or subgroup	Acupuncture	cture Standard dose antipsychotics		Risk Ratio				Weight Risk Ratio	
	n/N	n/N	_	M-H	, Fixed,	95% CI		M-H, Fixed, 95% CI	
Total (95% CI)	270	151						Not estimat	ole
Total events: 0 (Acupuncture), 0 (	Standard dose antipsy	chotics)							
Heterogeneity: Not applicable									
Test for overall effect: Not applica	ble								
	Fa	vours Acupuncture	0.01	0.1	1	10	100	Favours Standard dose antipsychotics	

### Analysis 3.6. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 6 Adverse effects: 1. General - average scores (TESS, endpoint, short-term) - Skewed data.

Adverse effects: 1. General - average scores (TESS, endpoint, short-term) - Skewed data											
Study	Intervention Mean SD N Note										
traditional - Wang 2006	Traditional acupuncture	2.43	3.17	16	P < 0.01						
traditional - Wang 2006	Antipsychotics	6.93	5.28	16							

### Analysis 3.7. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 7 Adverse effects: 2. Specific - extrapyramidal symptoms (short-term).

Study or subgroup	Acupuncture	Standard dose antipsychotics		Risk Ratio				Weight	Risk Ratio
	n/N	n/N		M-H, Fixe	ed, 95%	6 CI			M-H, Fixed, 95% Cl
laser - Zhang 1991	0/11	8/10	←					100%	0.05[0,0.83]
Total (95% CI)	11	10						100%	0.05[0,0.83]
Total events: 0 (Acupuncture), 8 (Sta	ndard dose antipsyc	hotics)							
Heterogeneity: Not applicable									
Test for overall effect: Z=2.09(P=0.04	)								
	Fav	ours Acupuncture	0.01	0.1	1	10	100	Favours Standard dos	se antipsychotics

# Analysis 3.8. Comparison 3 ACUPUNCTURE versus STANDARD DOSE ANTIPSYCHOTICS, Outcome 8 Adverse effects: 3. Specific - numbness over the ear, upper extremity and chest on the treated side (short-term).

Study or subgroup	Acupuncture	Standard dose antipsychotics	tandard dose ntipsychotics		Risk Ratio			Weight	Risk Ratio
	n/N	n/N		М-Н, А	ixed, 95% C	:1			M-H, Fixed, 95% CI
laser - Liu 1986	1/20	0/20						100%	3[0.13,69.52]
Total (95% CI)	20	20						100%	3[0.13,69.52]
Total events: 1 (Acupuncture), 0 (Sta	ndard dose antipsyc	hotics)							
Heterogeneity: Not applicable									
Test for overall effect: Z=0.69(P=0.49)									
	Fav	ours Acupuncture	0.01	0.1	1	10	100	Favours Standard dos	se antipsychotics

#### Comparison 4. ACUPUNCTURE added to TCM DRUG versus TCM DRUG

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Global state: Not improved, endpoint (short-term)	3		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
1.1 talk, behaviour and expression still ex- isted as before + reduction of BPRS < 29%	2	360	Risk Ratio (M-H, Fixed, 95% CI)	0.11 [0.02, 0.59]
1.2 no change in symptoms	1	94	Risk Ratio (M-H, Fixed, 95% CI)	0.77 [0.57, 1.06]
2 Behaviour: Leaving the study early (short-term)	3	454	Risk Ratio (M-H, Fixed, 95% CI)	0.0 [0.0, 0.0]

# Analysis 4.1. Comparison 4 ACUPUNCTURE added to TCM DRUG versus TCM DRUG, Outcome 1 Global state: Not improved, endpoint (short-term).

Study or subgroup	Acupunc- ture added to TCM drug	TCM drug	Risk Ratio		Weight	Risk Ratio
	n/N	n/N	M-H, Fixed, 959	% CI		M-H, Fixed, 95% CI
4.1.1 talk, behaviour and expression of BPRS < 29%	n still existed as be	fore + reduction				
traditional - Zhao 2005a	1/90	6/90			44.44%	0.17[0.02,1.36]
traditional - Zhao 2005b	0/90	7/90			55.56%	0.07[0,1.15]
Subtotal (95% CI)	180	180			100%	0.11[0.02,0.59]
Total events: 1 (Acupuncture added to	o TCM drug), 13 (TCM	l drug)				
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.27, df=	1(P=0.61); I <sup>2</sup> =0%					
Test for overall effect: Z=2.57(P=0.01)						
4.1.2 no change in symptoms						
electro - Zhang 1987	27/49	32/45	<del>-+</del> -		100%	0.77[0.57,1.06]
Subtotal (95% CI)	49	45	•		100%	0.77[0.57,1.06]
Total events: 27 (Acupuncture added	to TCM drug), 32 (TC	M drug)				
Heterogeneity: Not applicable						
Test for overall effect: Z=1.59(P=0.11)						
Test for subgroup differences: Chi <sup>2</sup> =4.	99, df=1 (P=0.03), I <sup>2</sup> =	79.96%				
Fav	ours Acupuncture ad	ded to TCM drug	0.005 0.1 1	10 200 Fa	vours TCM drug	

# Analysis 4.2. Comparison 4 ACUPUNCTURE added to TCM DRUG versus TCM DRUG, Outcome 2 Behaviour: Leaving the study early (short-term).

Study or subgroup	Acupunc- ture added to TCM drug	TCM drug		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		М-Н,	Fixed, 95	% CI			M-H, Fixed, 95% CI
electro - Zhang 1987	0/49	0/45							Not estimable
traditional - Zhao 2005a	0/90	0/90							Not estimable
traditional - Zhao 2005b	0/90	0/90							Not estimable
	Favours Acupuncture ad	ded to TCM drug	0.01	0.1	1	10	100	Favours TCM drug	

Acupuncture for schizophrenia (Review)



Study or subgroup	Acupunc- ture added to TCM drug	TCM drug		Risk Ratio			Weight	Risk Ratio	
	n/N	n/N		M-H	, Fixed, 95	5% CI			M-H, Fixed, 95% CI
Total (95% CI)	229	225							Not estimable
Total events: 0 (Acupuncture add	led to TCM drug), 0 (TCM	drug)							
Heterogeneity: Not applicable									
Test for overall effect: Not application	able								
	Favours Acupuncture a	dded to TCM drug	0.01	0.1	1	10	100	Favours TCM drug	

### Comparison 5. ACUPUNCTURE versus TCM DRUG

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Global state: Not improved, endpoint (short-term)	3		Risk Ratio (M-H, Fixed, 95% CI)	Subtotals only
1.1 talk, behaviour and expression still ex- isted as before + reduction of BPRS < 29%	2	360	Risk Ratio (M-H, Fixed, 95% CI)	1.46 [0.74, 2.87]
1.2 no change in symptoms	1	88	Risk Ratio (M-H, Fixed, 95% CI)	0.52 [0.34, 0.80]
2 Behaviour: Leaving the study early (short-term)	3	328	Risk Ratio (M-H, Fixed, 95% CI)	0.0 [0.0, 0.0]

### Analysis 5.1. Comparison 5 ACUPUNCTURE versus TCM DRUG, Outcome 1 Global state: Not improved, endpoint (short-term).

Study or subgroup	Acupuncture	TCM drug		Risk Ratio		Weight	<b>Risk Ratio</b>
	n/N	n/N	M-I	l, Fixed, 95% Cl			M-H, Fixed, 95% Cl
5.1.1 talk, behaviour and expression of BPRS < 29%	on still existed as be	fore + reduction					
traditional - Zhao 2005a	10/90	6/90				46.15%	1.67[0.63,4.39]
traditional - Zhao 2005b	9/90	7/90		<b></b>		53.85%	1.29[0.5,3.3]
Subtotal (95% CI)	180	180		-		100%	1.46[0.74,2.87]
Total events: 19 (Acupuncture), 13 (7	CM drug)						
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.14, df	=1(P=0.71); I <sup>2</sup> =0%						
Test for overall effect: Z=1.1(P=0.27)							
5.1.2 no change in symptoms							
electro - Zhang 1987	16/43	32/45		<b></b>		100%	0.52[0.34,0.8]
Subtotal (95% CI)	43	45		$\overline{\bullet}$		100%	0.52[0.34,0.8]
Total events: 16 (Acupuncture), 32 (1	CM drug)						
Heterogeneity: Not applicable							
Test for overall effect: Z=2.95(P=0)							
Test for subgroup differences: Chi <sup>2</sup> =6	5.33, df=1 (P=0.01), I <sup>2</sup> =	=84.19%			1		
	Favo	ours Acupuncture	0.01 0.1	1 10	100	Favours TCM drug	



### Analysis 5.2. Comparison 5 ACUPUNCTURE versus TCM DRUG, Outcome 2 Behaviour: Leaving the study early (short-term).

Study or subgroup	Acupuncture	TCM drug			Risk Ratio			Weight	<b>Risk Ratio</b>
	n/N	n/N		M-H	l, Fixed, 95	% CI			M-H, Fixed, 95% Cl
electro - Zhang 1987	0/43	0/45							Not estimable
traditional - Zhao 2005a	0/90	0/30							Not estimable
traditional - Zhao 2005b	0/90	0/30							Not estimable
Total (95% CI)	223	105							Not estimable
Total events: 0 (Acupuncture), 0 (TCM	l drug)								
Heterogeneity: Not applicable									
Test for overall effect: Not applicable									
	Favo	urs Acupuncture	0.01	0.1	1	10	100	Favours TCM drug	

### Comparison 6. ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Behaviour: Leaving the study early (short-term)	1	68	Risk Ratio (M-H, Fixed, 95% CI)	0.0 [0.0, 0.0]
2 Adverse effects: 1. Specific - back pain (short-term)	1	68	Risk Ratio (M-H, Fixed, 95% CI)	0.67 [0.12, 3.74]
3 Adverse effects: 2. Spinal fracture (short-term)	1	68	Risk Ratio (M-H, Fixed, 95% CI)	0.33 [0.14, 0.81]

# Analysis 6.1. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 1 Behaviour: Leaving the study early (short-term).

Study or subgroup	Electric acupunc- ture convul- sive therapy	Electroconvul- sive therapy			Risk Ratio	Ratio		Weight	Risk Ratio
	n/N	n/N		M-H	l, Fixed, 95%	% CI			M-H, Fixed, 95% CI
EACT - Xue 1987	0/34	0/34							Not estimable
Total (95% CI)	34	34							Not estimable
Total events: 0 (Electric acupuncture vulsive therapy)	convulsive therapy	), 0 (Electrocon-							
Heterogeneity: Not applicable									
Test for overall effect: Not applicable									
Favours Ele	ectric acupuncture of	convulsive therapy	0.01	0.1	1	10	100	Favours Electroconvu	lsive therapy

# Analysis 6.2. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 2 Adverse effects: 1. Specific - back pain (short-term).

Study or subgroup	Electric acupunc- ture convul- sive therapy	Electroconvul- sive therapy			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H	, Fixed, 95%	CI			M-H, Fixed, 95% Cl
EACT - Xue 1987	2/34	3/34						100%	0.67[0.12,3.74]
Total (95% CI)	34	34						100%	0.67[0.12,3.74]
Total events: 2 (Electric acupuncture vulsive therapy)	convulsive therapy)	, 3 (Electrocon-							
Heterogeneity: Not applicable									
Test for overall effect: Z=0.46(P=0.64)				1		1			
Favours Ele	ectric acupuncture c	onvulsive therapy	0.01	0.1	1	10	100	Favours Electroconvu	lsive therapy

# Analysis 6.3. Comparison 6 ELECTRIC ACUPUNCTURE CONVULSIVE THERAPY versus ELECTROCONVULSIVE THERAPY, Outcome 3 Adverse effects: 2. Spinal fracture (short-term).

Study or subgroup	Electric acupunc- ture convul- sive therapy	Electroconvul- sive therapy			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-H	, Fixed, 95%	6 CI			M-H, Fixed, 95% Cl
EACT - Xue 1987	5/34	15/34		-				100%	0.33[0.14,0.81]
Total (95% CI)	34	34						100%	0.33[0.14,0.81]
Total events: 5 (Electric acupuncture vulsive therapy)	convulsive therapy	), 15 (Electrocon-							
Heterogeneity: Not applicable									
Test for overall effect: Z=2.41(P=0.02)									
Favours Ele	ectric acupuncture	convulsive therapy	0.01	0.1	1	10	100	Favours Electroconvu	Ilsive therapy

#### ADDITIONAL TABLES

#### Table 1. Other outcomes authors reported

#### **Global state: Adding medication**

Study	Drug	Events/Acupunc- ture group	Events/Control group
		(n/N)	(n/N)
electro - Cui 2000	Artane	6/30	11/30
	Promethazine	3/30	6/30
	Propranolol	2/30	4/30
	Chlorpheniramine	0/30	1/30

Acupuncture for schizophrenia (Review)



### Table 1. Other outcomes authors reported (Continued)

electro - Wang 2005	Propranolol	7/40	7/35
	Artane	6/40	9/35
	Benzodiazepine drugs	4/40	7/35
Mental state: Time t	o auditory hallucinations disanneared (days)		
Mental States Time (	o dualtory naturellations disappeared (days)		
Study	Intervention	Mean	SD
Study	Intervention Laser acupuncture added to standard dose antipsychotics	<b>Mean</b> 19	<b>SD</b> 4.1

### Table 2. Adverse effects of acupuncture

Intervention	Term	Adverse effects		Adverse ef- fects/In- tervention group	Study
				(n/N)	
Traditional acupuncture					
Traditional acupuncture added to standard dose	Short term	Extrapyramidal symp- toms	Overall	1/30	traditional - Ma 2008
standard dose antipsy- chotics		Anticholinergic symp- toms	Dry mouth	3/30	_
		Cardiovascular	Dizziness	3/30	_
			Tachycardia	2/30	_
	Medium term	Extrapyramidal symp- toms	Overall	5/47	traditional - Liu 2010
		Central Nervous Sys-	Anxiety	2/47	_
		tem	Insomnia	3/47	_
		Cardiovascular symp- toms (or headache)	Dizziness or headache	1/47	_
		Gastrointestinal sys- tem	Unspecifid gastroin- testinal symptoms	2/47	
		Lab test	Liver function abnor- mal	2/47	

#### Electroacupuncture

### Table 2. Adverse effects of acupuncture (Continued)

Electroacupuncture (or added to TCM drug) versus TCM drug	Short term	Hold breath, facial cyanosis, arrhythmia, tran- sient increase of blood pressure, injury of teeth, tongue and lips, epileptic attacks with strong stimulation		not reported	electro - Zhang 1987
Electroacupuncture added to standard dose antipsy- chotics versus standard	Short term	rm Extrapyramidal symp- toms	Overall	18/73	electro - Chen 2006; electro - Wang 2005
dose anapsychotics			Tremor	5/40	electro - Wang – 2005
			Akathisia	6/40	
		Central Nervous Sys- tem	Insomnia	4/73	electro - Chen 2006; electro - Wang 2005
		Anticholinergic symp-	Dry mouth	6/73	
		toms	Blurred vision	4/73	
			Constipation	3/73	
			Nausea and vomiting	3/73	_
		Cardiovascular symp-	Dizziness	3/73	
			Tachycardia	10/78	electro - Chen 2008; electro - Yao 2006
		Metabolic system	Weight gain	3/73	electro - Chen 2006; electro - Wang 2005
		Lab test	Liver function abnor- mal	4/118	electro - Chen 2006; electro - Wang 2005
			ECG abnormal (my- ocardial ischaemia)	1/118	electro - Yao 2006
			blood routine test ab- normal (leukocyte change)	4/118	
	Medium term	Extrapyramidal symp- toms	Overall	6/30	electro - Chen — 2008 —
			Myotonia	1/30	
			Tremor	2/30	
			Akathisia	3/30	
		Central Nervous Sys-	Insomnia	1/30	_
			Headache	2/30	_

### Table 2. Adverse effects of acupuncture (Continued)

		Anticholinergic symp- toms	Blurred vision	1/30	
		toms	Sweating	3/30	-
Electroacupuncture added low dose antipsychotics	Short term	Extrapyramidal symp- toms	Overall	5/30	electro - Cui 2000
tipsychotics		Central Nervous Sys- tem	Sleepiness	1/30	-
		Anticholinergic symp-	Dry mouth	4/30	
			Blurred vision	2/30	
		Cardiovascular symp- toms	Tachycardia	5/30	-
Laser acupuncture					
Laser acupuncture ver- sus standard dose antipsy-	Short term	Numbness over the ear, chest on the treated side	upper extremity and e	1/20	laser - Liu 1986
		Sensation of heat at irra of plugged auditory can	diated site and a feeling al	one third of participants of He-Ne laser irradiation group	
Laser acupuncture added to low dose antipsychotics versus standard dose an- tipsychotics	Short term	Extrapyramidal symp- toms	Overall	7/10	laser - Zhang 1991
Acupoint catgut treatment					
Acupoint catgut treat- ment added to standard dose antipsychotics ver- sus standard dose antipsy- chotics	Short term	Local pain when eating		not reported	acupoint cat - Wang 1997
Acupoint catgut treatment	Short term	Cardiovascular symp-	Dizziness	2/88	acupoint cat -
tipsychotics versus stan- dard dose antipsychotics		toms	Tachycardia	2/88	- 30112005
Electric acupuncture convu	Isive therapy				
Electric acupuncture con-	Short term	Back pain		2/34	EACT - Xue 1987
electroconvulsive therapy		Spinal fracture		5/34	-

### Table 3. Summary of acupuncture methods

Study	Kind of acupuncture	Acupoints choice	Fequency and duration	Sham acupunc- ture (yes/no)
traditional - Bouhlel 2011	Traditional acupuncture	Local points, distal points di- agnosed by TCM	Once for 20 minutes and 3 times a week; total 10 times	Yes
traditional - Liu 2010	-	Main acupoints and adjunct acupoints according to the TCM syndrome differentia- tion	Manipulated needles every 10 minutes; for 30 minutes; 4 to 5 times a week and no less than 4 times; 1 month as 1 treatment course, 3 treatment course	No
traditional - Luo 2006	-	Two acupoint groups in turn	For 30 minutes; once a day; 20 days as a treatment course and began anoth- er treatment course after 10 non-treat- ment days; 3 months treatment cours- es	No
traditional - Ma 2008	-	Juque, Tanzhong, Taichong, Jianshi, Fenglong, Daling, Yingtang	For 30 minutes; once a day; 5 days treatment with 2 days non-treatment interval; 6 weeks	No
traditional - Tang 2005	-	According to the type of TCM	Manipulated needles every 10 minutes; for 30 minutes; 3 to 4 times a week and not less than 3 times; 3 weeks as a treatment course and 1 week interval; total 3 treatment courses	No
traditional - Wang 2006	-	Taichong (reducing method), Hegu (reducing method), Neiguan (straight insert- ed), Daling (straight insert- ed), Renzhong (reducing method), Dazhui (straight inserted) + other acupoints choice according to type of TCM	Continued to manipulate needles 3 to 5 minutes every ten minutes; for 45 minutes; once a day; 15 times as a treatment course; 2 treatment courses	No
traditional - Xu 2004	-	Main acupoints and adjunct acupoints (according to the TCM type of schizophrenia) and special acupoints (ac- cording to symptoms)	Manipulated needles about 3 minutes and once ten minutes; total for about 30 minutes; at first once a day then re- duced to once every two days when patient was in stable condition and once a week after psychiatric symp- toms disappeared; 80 days	No
traditional - Zhao 2005a	-	Shuigou, Shaoshang, Yingbai, Fengfu, Daling, Quchi, Feng- long	Once a day; for 30 minutes; 60 days	No
traditional - Zhao 2005b	-	Xinshu, Ganshu, Pishu, Shen- men, Fenglong	Once a day; for 30 minutes; 60 days	No
electro - Chen 2006	Electroacupunc- ture	Baihui and Yingtang	For 50 minutes, once a day, 6 weeks	No
electro - Chen 2008	-	Two acupoints groups in turn - Baihui and Neiguan (dou-	For 45 minutes; once a day; five days a week (except Saterday and Sunday); 12 weeks as a treatment course	No

Acupuncture for schizophrenia (Review)



	ble) group or Shuigou and Sanyingjiao (double) group		
electro - Cheng 2009	Tinggong, Tinghui, Yifeng, Daling, Neiguan and Sanyi- jiao	For 20 minutes; 5 times a week; total 30 times (6 weeks)	Yes; needles were inserted less than 5 mm superficially and about 20 mm away from each corresponding acupoint and electroacupunc- ture connected was with no elec- trical current for 20 minutes
electro - Cui 2000	According to different symp- toms	For 30 minutes; once two days; 20 times as a treatment course	No
electro - Ding 2005	Tanzhong, Zhongwan, Shen- men, Fenglong (double), Tai- chong (double), Neiguan (double)	For 5 minutes; once two days	No
electro - Wang 2005	Baihui, Shangxing, Ying- tang, Sanyingjiao (double), Neiguan (double); added Zhongwan and Zusanli when with gastrointestinal un- comfortable symptoms and added Fenglong when with symptoms of the type of stagnation of phlegm and Qi	For 45 minutes; 5 times a week; 20 times as a treatment course; 8 weeks	No
electro - Xiong 2010	Baihui and Taiyang (double)	Firstly continued to stimulate strong- ly 3 to 4 seconds then reduced the strength and frequency quickly, after patient's breath and face returned to normal and continued 30 to 60 sec- onds stimulated again; continued to stimulate 8 to 10 times; 3 times a week, total 8 weeks	No
electro - Yao 2006	Baihui, Fenglong, Houxi, Gan- shu	For 30 minutes; once a day; 20 to 30 times as a treatment course; total 2 treatment courses	No
electro - Zhang 1987	Yifeng, Tinggong, Tounjie, Chengling, Linqi, Baihui, Dingshen	Twice a day; adjustment of treatment depended upon the condition of the patient; 20 days	No
electro - Zhang 1993	Group one [Yingtang and Bai- hui] and group two [Shenting and Yamen]	For 45 minutes; once a day; 8 weeks as a treatment course	No
electro - Zhang 2001	Baihui and Yingtang	For 45 minutes; once a day; 5 days a week except Saturday and Sunday; 6 weeks as a treatment course	No

Acupuncture for schizophrenia (Review)

=

electro - Zhou 1997	, <b>,</b>	According to TCM types; main acupoints (Yintang tou xin- qu, Daling, Neiguan, Taiyang) and supplemental acupoints (Zusanli [Yang deficiency]	Once a day except Sunday; 36 times as a treatment course (6 weeks)	No
acupoint inj - Pan 2002	Acupoint injec- tion	According to the type of TCM (with Salviae Miltiorrhizae)	Two side acupoints in turn; once a day; 10 days as a treatment course; 7 days interval between two courses; 3 treat- ment courses	No
acupoint inj - Wang 2000	-	Tinggong (double) (with Clonazepam)	Once two days; 7 times	No
acupoint inj - Yang 2000	_	Tinggong (double) (Sulpiride)	Once a day; 5 times as a treatment course; 2 intermittent treatment cours- es each month; total half a year	No
laser - Liu 1986	Laser acupunc- ture	Ermen (double) (He-Ne laser irradiation)	For 15 minutes; once a day except Sun- day; 30 times as a treatment course	Yes; sham laser irradiation; the tube of the laser emitter pointed in the direction of the ear with- out real irradia- tion
laser - Ma 1999	-	Tinggong and Tinghui (He-Ne laser irradiation)	For 30 minutes; once a day except Sun- day	No
laser - Zhang 1991	_	Two acupoint groups were used every other day alter- nately - group 1 (Dazhui and Shenting); group 2 (Taiyang [double])	For 15 minutes; once a day (except Sunday); 5 weeks as a treatment course	Yes; needles fixed with tape on acupoints
acupoint cat - Sun 2005	Acupoint catgut treatment	Tinggong	Once 7-10 days; 6 weeks as a treatment course	No
acupoint cat - Wang 1997	-	Tinggong (double)	10 days as a treatment course; total 3 treatment courses	No
EACT - Xue 1987	Electric acupunc- ture convulsive therapy	Renzhong and Baihui	Once every two days; 12 times as a treatment course	No

### Table 3. Summary of acupuncture methods (Continued)

### Table 4. Missing outcomes of each comparison

Comparison	Missing outcomes
ACUPUNCTURE added to STANDARD DOSE ANTIPSYCHOTICS versus STAN- DARD DOSE ANTIPSYCHOTICS	Death, engagement with services, satisfaction with treatment, quality of life, economic outcomes


# Table 4. Missing outcomes of each comparison (Continued)

ACUPUNCTURE added to LOW DOSE AN- TIPSYCHOTICS versus STANDARD DOSE ANTIPSYCHOTICS	Death, service outcomes, engagement with services, satisfaction with treatment, quality of life, economic outcomes
ACUPUNCTURE versus STANDARD AN- TIPSYCHOTICS	Death, service outcomes, engagement with services, satisfaction with treatment, quality of life, economic outcomes
ACUPUNCTURE added to TCM DRUG ver- sus TCM DRUG	Death, mental state, service outcomes, adverse effects, engagement with services, satis- faction with treatment, quality of life, economic outcomes
ACUPUNCURE versus TCM DRUG	Death, mental state, service outcomes, adverse effects, engagement with services, satis- faction with treatment, quality of life, economic outcomes
ELECTRIC ACUPUCNTURE CONVULSIVE THERAPY versus ELECTRIC ACUPUNC- TURE CONVULSIVE THERAPY	Death, global state, mental state, service outcomes, engagement with services, satisfac- tion with treatment, quality of life, economic outcomes

## Table 5. Other comparisons of relevance to this systematic review

Comparison	Trial
Different acupoints	Ma 2002; electro - Zhang 1987
Acupoints vs non-acupoints	laser - Liu 1986

# Table 6. Suggested design for a trial Methods Allocation: randomised, clearly described, concealed. Blindness: double, described and tested. Duration: 4 week washout period + 24 weeks treatment period. Follow-up: 2 years. **Participants** Diagnosis: schizophrenia (DSM V) with one TCM type according to TCM diagnosis standard. $N = 300^*$ . Age: any. Sex: both. History: duration of schizophrenia over 1 years; never receive acupuncture previously\*\* (at least). Interventions 1. Electroacupuncture: N = 150. 2. Sham electroacupuncture: N = 150. The acupoints and relative parameters clearly described. Outcomes 1. Death 2. Global state 3. Mental state 4. Behaviour 5. Service outcomes 6. Adverse effects

Acupuncture for schizophrenia (Review)

Copyright  $\odot$  2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

# Table 6. Suggested design for a trial (Continued)

Notes	<ul> <li>* Powered to be able to identify a difference of ~20% between groups for primary outcome with ad-equate degree of certainty.</li> <li>** This is to maximise blinding.</li> </ul>	
	10. Economic outcome	
	9. Quality of life	
	8. Satisfaction with treatment	
	7. Engagement with service	

# APPENDICES

# Appendix 1. Previous type of interventions (2005)

1. Acupuncture/electro-acupuncture with or without moxibustion (a Traditional Chinese Medicine technique that involves the burning of mugwort to facilitate healing) or laser treatment, administered solely or in conjunction with antipsychotic drugs.

2. Placebo (sham acupuncture) or no treatment.

3. Antipsychotic drugs produced by pharmaceutical companies: any compound, dose, pattern or means of administration.

# Appendix 2. Previous search strategy (2005)

## 1. Electronic searches

We searched the Cochrane Schizophrenia Group's Trials Register (April 2005) using the phrase: [(\*acup\* OR \*moxibustion\*) in REFERENCE and (\*acupuncture\* OR \*moxibustion\*) in STUDY]

This register is compiled by systematic searches of major databases (Biological Abstracts CINAHL, *The Cochrane Library*, EMBASE, MEDLINE, RUSSMED, LILACS, PSYNDEX and PsycLIT) hand searches and conference proceedings (see Group Module).

2. We also inspected the references of all identified studies (included and excluded) for further relevant trials.

# Appendix 3. Previous data collection and analysis methods (2005)

# 1. Selection of studies

We (JR, JX) inspected all reports of studies identified from the search. A randomly selected sample of 10% of all the reports were reinspected in order to ensure that the selection was reliable. Where disagreement occurred, we resolved this by discussion and if there was still doubt we acquired the full article for further inspection. Once the full articles were obtained, we independently decided whether or not they met the review criteria. Again, where disagreement occurred, we resolved this by discussion and when this was not possible we sought further information from first authors and added these trials to the list of those awaiting assessment.

# 2. Assessment of methodological quality

We allocated trials to three quality categories, as described in the Cochrane Collaboration Handbook (Alderson 2004). When disputes arose as to which category a trial was allocated, again, we attempted resolution by discussion. When this was not possible and further information was necessary to clarify into which category to allocate the trial, we did not enter data but allocated the trial to the list of those awaiting assessment. We only included trials in Category A or B in the review.

A. Low risk of bias (adequate allocation concealment)

B. Moderate risk of bias (some doubt about the results)

C. High risk of bias (inadequate allocation concealment). For the purpose of the analysis in this review, trials were included if they met the Cochrane Handbook criteria A or B.

## 3. Data extraction

3.1 Reliable extraction

We independently extracted data from selected trials. When disputes arose, we attempted resolution by discussion. When this was not possible and further information was necessary to resolve the dilemma, we did not enter data but added this outcome of the trial to the list of those awaiting assessment.

# 3.2 Intention to treat analysis



We excluded data from studies where more than 50% of participants in any group were lost to follow-up, except for the outcome of 'leaving the study early'. In studies with less than 50% dropout rate, everyone allocated to the intervention was counted whether or not they completed follow-up. We considered those leaving early to have had the negative outcome, except for the event of death and adverse effects.

Where attrition rates were high (25% to 50%), we analysed the impact of including this type of data in a sensitivity analysis. If inclusion of high attrition data resulted in a substantive change in the estimate of effect, then we did not pool this data, but presented the data separately.

## 4. Data analysis

## 4.1 Dichotomous/binary data

In the review we used relative risk (RR), (fixed-effect) and 95% confidence interval (CI). Where possible, we made efforts to convert relevant categorical or continuous outcome measures to dichotomous data by identifying cut off points on rating scales and dividing them into groups accordingly i.e.. 'moderate or severe impairment' and 'no better or worse'.

4.1.1 Summary statistic: For binary outcomes we calculated a standard estimate of the relative risk (RR) and its 95% confidence intervals (CI) (fixed-effect). Where possible, we estimated the number needed to treat (NNT) using an on-line calculator (http://www.nntonline.net/). If heterogeneity was found (see section 5) we used a random-effects model.

## 4.2 Continuous data

4.2.1 Skewed data: continuous data on clinical and social outcomes are often not normally distributed. To avoid the pitfall of applying parametric tests to non-parametric data, the following standards are applied to all data before inclusion: (a) standard deviations and means were reported in the paper or were obtainable from the authors; (b) when a scale started from the finite number zero, the standard deviation, when multiplied by two, was less than the mean (as otherwise the mean was unlikely to be an appropriate measure of the centre of the distribution, (Altman 1996); (c) if a scale started from a positive value (such as PANSS which can have values from 30-210) the calculation described above in (b) was modified to take the scale starting point into account. In these cases skewness is present if 2SD>(S-Smin), where S is the mean score and Smin is the minimum score. Endpoint scores on scales often have a finite start and end point and these rules can be applied to them.

4.2.2 Summary statistic: For continuous outcomes we calculated weighted mean differences (WMD) and respective 95% CI (fixed-effect). If heterogeneity was found (see section 5) we used a random-effects model.

4.2.3 Valid scales: A wide range of instruments are available to measure mental health outcomes. These instruments vary in quality and it has been shown that the use of rating scales which have not been described in a peer-reviewed journal (Marshall 2000) are associated with bias, or may not be valid, or even ad hoc. Therefore, some minimum standards were set: (a) the psychometric properties of the instrument should have been described in a peer-reviewed journal; (b) the instrument should either be a self-report, or completed by an independent rater or relative (not the therapist); and (c) the instrument should be a global assessment of an area of functioning.

4.2.4 Endpoint versus change data: where possible, we presented endpoint data and if both endpoint and change data were available for the same outcomes then we only reported the former in this review.

### 4.2.5 Cluster trials

Studies increasingly employ 'cluster randomisation' (such as randomisation by clinician or practice) but analysis and pooling of clustered data poses problems. Firstly, authors often fail to account for intra class correlation in clustered studies, leading to a 'unit of analysis' error (Divine 1992) whereby P values are spuriously low, confidence intervals unduly narrow and statistical significance overestimated. This causes type I errors (Bland 1997, Gulliford 1999).

Where clustering was not accounted for in primary studies, we presented the data in a table, with a (\*) symbol to indicate the presence of a probable unit of analysis error. In subsequent versions of this review we will seek to contact first authors of studies to obtain intra-class correlation co-efficients of their clustered data and to adjust for this using accepted methods (Gulliford 1999). Where clustering has been incorporated into the analysis of primary studies, we will also present these data as if from a non-cluster randomised study, but adjusted for the clustering effect.

We have sought statistical advice and have been advised that the binary data as presented in a report should be divided by a 'design effect'. This is calculated using the mean number of participants per cluster (m) and the intraclass correlation co-efficient (ICC) [Design effect = 1+(m-1)\*ICC] (Donner 2002). If the ICC was not reported it was assumed to be 0.1 (Ukoumunne 1999).

#### 5. Test for heterogeneity

Firstly, we considered all of the included studies within any comparison to judge clinical heterogeneity. Then we visually inspected graphs used to investigate the possibility of statistical heterogeneity and supplemented this by using, primarily, the I-squared statistic. This provides an estimate of the percentage of variability due to heterogeneity rather than chance alone. Where the I-squared estimate was greater than or equal to 75%, we interpreted this as indicating the presence of high levels of heterogeneity (Higgins 2003). If inconsistency was high, we did not summate the data, but presented it separately and reasons for heterogeneity were investigated.



# 6. Addressing publication bias

We entered all data from the included studies into a funnel graph (trial effect against trial size) in an attempt to investigate the likelihood of overt publication bias (Egger 1997).

## **Appendix 4. Previous description of studies**

#### 1. Excluded studies

We excluded three studies. Xue 1985 and Zhuge 1993 were not randomised. Zhong 1995 compared electro-acupuncture to computerised electrode.

### 2. Awaiting assessment

No studies are awaiting assessment.

#### 3. Ongoing studies

We are not aware of any ongoing studies.

#### 4. Included studies

We included five studies all of which were randomised. Only electro - Zhou 1997 reported on blinding, in which the raters were blind to treatment allocation.

#### 4.1 Length of trials

All included studies were short-term (less than three months). The shortest study electro - Zhang 1987 lasted for 20 days. laser - Zhang 1991 was for five weeks, electro - Zhou 1997 and electro - Zhang 2001 were six week studies and Zhang 1994 was the longest study lasting eight weeks.

#### 4.2 Participants

All participants were diagnosed with schizophrenia. electro - Zhou 1997, laser - Zhang 1991 and electro - Zhang 2001 used operationalised criteria (DSM/ICD/CCMD). electro - Zhang 1987, Zhang 1994 did not report using predefined diagnostic criteria. electro - Zhou 1997, electro - Zhang 1987, laser - Zhang 1987, laser - Zhang 1991 and electro - Zhang 2001 used male and female participants, whilst Zhang 1994 did not report on gender.

## 4.3 Setting

All five trials were undertaken in a hospital setting.

#### 4.4 Study size

The numbers of participants randomised ranged between 31 laser - Zhang 1991 to 88 electro - Zhang 1987. Forty people were randomised by electro - Zhou 1997, 69 by Zhang 1994 and 42 by electro - Zhang 2001.

#### 4.5 Interventions

The treatment groups received acupuncture exclusively or in combination with antipsychotics. All the comparator groups received antipsychotics. electro - Zhou 1997 used electro-acupuncture in combination with reduced dose antipsychotics versus chlorpromazine equivalents (~560mg/day). The treatment group in electro - Zhang 1987 received electro-acupuncture and the comparator group were given chlorpromazine (300-600 mg/day). laser - Zhang 1991 randomised people into three comparator groups, laser-acupuncture with moxibustion, laser acupuncture with moxibustion and reduced dose chlorpromazine (150-300 mg/day), or chlorpromazine (350-600 mg/day). Zhang 1994 compared electro-acupuncture to chlorpromazine equivalents (~458 mg/day). electro - Zhang 2001 compared electro-acupuncture combined with antipsychotics to antipsychotics alone but did not report on the acupoints used or dosages. Details of acupuncture points, needles and electro-acupuncture frequencies used (when reported) are listed in the included studies table.

## 4.6 Outcomes

All data outcomes were reported as short-term (less than three months). Data from some studies were unusable because raw scores were not presented. Instead outcomes were reported as P values without means and standard deviations being provided, or the scale used was not identified. electro - Zhang 2001 predefined no clinical improvement as HAMD scores reduced by </= 25%. electro - Zhou 1997 predefined no clinical improvement as BPRS scores reduced by </= 20%.

4.6.1 Outcome scales: details of scales that provided usable data are shown below. Reasons for exclusion of data from instruments are given under 'outcomes' in the included studies table.

#### 4.6.1.1 Global state scales

#### 4.6.1.1.1 Clinical Global Impression Scale - CGI (Guy 1970)

The CGI is a three-item scale commonly used in studies on schizophrenia that enables clinicians to quantify severity of illness and overall clinical improvement. The items are: severity of illness, global improvement and efficacy index. A seven-point scoring system is usually used with low scores indicating decreased severity and/or greater recovery. electro - Zhou 1997 and laser - Zhang 1991 reported CGI data.

#### 4.6.1.2 Mental state

4.6.1.2.1 Brief Psychiatric Rating Scale - BPRS (Overall 1962)

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd.

The BPRS is an 18-item scale measuring positive symptoms, general psychopathology and affective symptoms. The original scale has sixteen items, but a revised eighteen-item scale is commonly used. Scores can range from 0-126. Each item is rated on a seven-point scale, with high scores indicating more severe symptoms. electro - Zhou 1997, laser - Zhang 1991and Zhang 1994 all reported BPRS data.

#### 4.6.1.2.2 Scale for the Assessment of Negative Symptoms - SANS (Andreasen 1982)

This scale allows a global rating of the following negative symptoms: alogia (impoverished thinking), affective blunting, avolition-apathy, anhedonia-asociality and attention impairment. Assessments are made on a six-point scale (0 = not at all to 5 = severe). Higher scores indicate more symptoms. Data for this scale were reported by Zhang 1994.

## 4.6.1.2.3 Scale for the Assessment of Positive Symptoms - SAPS (Andreasen 1982)

This six-point scale gives a global rating of positive symptoms such as delusions, hallucinations and disordered thinking. Higher scores indicate more symptoms. Zhang 1994 was the only study to report SAPS data.

## 4.6.1.2.4 Hamilton Rating Scale for Depression - HAMD (Hamilton 1967)

This instrument is designed to be used only on patients already diagnosed as suffering from affective disorder of the depressive type. It is used for quantifying the results of an interview, and its value depends entirely on the skill of the interviewer in eliciting the necessary information. The scale contains 17 variables measured on either a five-point or a three-point rating scale, the latter being used where quantification of the variable is either difficult or impossible. Among the variables are: depressed mood, suicide, work and loss of interest, retardation, agitation, gastro-intestinal symptoms, general somatic symptoms, hypochondriasis, loss of insight, and loss of weight. It is useful to have two raters independently scoring a patient at the same interview. The scores of the patient are obtained by summing the scores of the two physicians. A score of 11 is generally regarded as indicative of a diagnosis of mild depression, 14-17 mild to moderate depression and >17 moderate to severe depression. electro - Zhang 2001 reported data for this scale.

## 4.6.1.2.5 Zung Depression Scale - ZDS (Zung 1965)

The Zung Self-Rating Depression Scale is a 20-item self-rated scale that is widely used as a screening tool, covering affective, psychological and somatic symptoms associated with depression. The questionnaire takes approximately ten minutes to complete and items are framed in terms of positive and negative statements. It can be effectively used in a variety of settings, including primary care, psychiatric clinics, drug trials and various research situations. Each item is scored on a Likert scale ranging from one to four. Most people with depression score between 50 and 69, while a score of 70 and above indicates severe depression. electro - Zhang 2001 reported data from this scale.

#### 4.6.1.3 Adverse events

## 4.6.1.3.1 Treatment Emergent Symptom Scale/Form - TESS/F (Guy 1976)

This checklist assesses a variety of characteristics for each adverse event, including severity, relationship to the drug, temporal characteristics (timing after a dose, duration and pattern during the day), contributing factors, course and action taken to counteract the effect. Symptoms can be listed a priori or can be recorded as observed by the investigator. Zhang 1994 reported data for this scale.

# **Appendix 5. Previous effects of interventions**

1. The search

The electronic search identified 18 reports. We were able to include five trials and we added three trials to the excluded studies table.

# 2. COMPARISON 1. ACUPUNCTURE versus ANTIPSYCHOTICS

#### 2.1. Global state

Two studies electro - Zhang 1987 and laser - Zhang 1991 reported on global state (not improved) with equivocal results.

## 2.2. Leaving the study early

There were no losses to follow-up in either group from two trials electro - Zhang 1987 and laser - Zhang 1991 by five weeks.

#### 2.3. Adverse effects

Extrapyramidal symptoms reported by laser - Zhang 1991 were lower in the acupuncture group with no participants experiencing this adverse effect. The control group had significantly higher numbers with eight out of ten people reported as having extrapyramidal symptoms (n = 21, RR 0.05 Cl 0.0 to 0.8, NNT 2 Cl 2 to 8).

# 3. COMPARISON 2. ACUPUNCTURE + ANTIPSYCHOTICS versus ANTIPSYCHOTICS

#### 3.1. Global state

One study laser - Zhang 1991 reported global state (not improved at five weeks) with equivocal results. Clinical Global Impression (severity of illness) scores were reported by electro - Zhou 1997 with equivocal scores. electro - Zhou 1997 also reported on Clinical Global Impression (global improvement) but data were skewed.

#### 3.2. Leaving the study early

Four trials (laser - Zhang 1991, Zhang 1994, electro - Zhou 1997, electro - Zhang 2001) reported as short-term outcomes that no participants left the study early (up to eight weeks).



# 3.3. Mental state

Dichotomised BPRS data were reported by electro - Zhou 1997 (not improved </ = 20% reduction at endpoint) with equivocal results. Continuous BPRS data (Zhang 1994, electro - Zhou 1997) were significant in favour of the acupuncture group (n = 109, WMD -4.31 CI -7.0 to -1.6). Mental state SANS and SAPS score were reported by Zhang 1994, but data were skewed and cannot be displayed graphically. Depression scores from the HAMD scale were reported by electro - Zhang 2001 with scores from this smaller study being significantly lower in the acupuncture group (n = 42, WMD -10.41 CI -12.8 to -8.0). When the HAMD scores were dichotomised to 'not improved', again, by electro - Zhang 2001 results significantly favoured the acupuncture group (n = 42, RR 0.17 CI 0.1 to 0.5, NNT 2 CI 2 to 3). electro - Zhang 2001 also reported on depression using the Zung Depression Scale with results at five weeks significantly favouring the acupuncture group (n = 42, WMD -24.25 CI -28.0 to -20.5).

## 3.4. Adverse effects

electro - Zhou 1997 reported TESS scores at six weeks, with results significantly favouring the acupuncture group (n = 40, WMD -0.50 CI -0.9 to -0.1). laser - Zhang 1991 found the incidences of extrapyramidal symptoms to be similar for both treatment groups.

## WHAT'S NEW

Date	Event	Description
19 August 2014	New citation required but conclusions have not changed	Results from new studies have not significantly changed overall conclusions from previous versions of this review.
17 January 2013	New search has been performed	New search carried out Feburary 2012, 30 studies now included in the review.

# HISTORY

Protocol first published: Issue 4, 2005 Review first published: Issue 4, 2005

Date	Event	Description
7 December 2011	Amended	Contact details updated.
10 November 2010	Amended	Contact details updated.
13 November 2008	Amended	CSG Data Extraction Tag added
23 April 2008	Amended	Converted to new review format.
18 May 2006	Amended	Minor update
5 July 2005	New citation required and conclusions have changed	Substantive amendment
28 June 2005	New citation required and conclusions have changed	Conclusions changed in light of update.
20 May 2005	New search has been performed	New studies found and included or excluded. Ammended in light of peer review comments.

# CONTRIBUTIONS OF AUTHORS

Xiaohon Shen - selected studies, extracted data and updated the full review.

Jun Xia - selected studies and extracted data.

Acupuncture for schizophrenia (Review) Copyright © 2014 The Cochrane Collaboration. Published by John Wiley & Sons, Ltd. Clive E Adams - assisted with data extraction, undertook analysis and writing the final report.

# DECLARATIONS OF INTEREST

Xiaohong Shen - none known.

Jun Xia - none known.

Clive E Adams - none known.

# SOURCES OF SUPPORT

# **Internal sources**

- Shanghai Shuguang Hospital affiliated to Shanghai University of Traditional Chinese Medicine, China.
- University of Nottingham, UK.

# **External sources**

• No sources of support supplied

# DIFFERENCES BETWEEN PROTOCOL AND REVIEW

# 1. Methodology update

The protocol used for this update has been changed and updated: taken from the most recent methodology of the Cochrane Schizophrenia Group, it acknowledges changes in Cochrane methodology since this review was first published. For example, use of 'Summary of findings' tables instead of calculating NNT/NNH and changes in the 'Risk of bias' tables and 'methods' being described in more detail. We do not think the use of these new methods affects the results or integrity of the review. For previous methodology please see Appendix 3.

# 2. Type of interventions

The title of this review is broad - 'Acupuncture for schizophrenia'. The previous version used a more narrow range of comparisons and, we felt, left out some that are important. We have added all categories of acupuncture alone or in combination regimens compared with placebo (or no treatment) or any other treatments. For previous types of interventions please see Appendix 1.

# INDEX TERMS

# **Medical Subject Headings (MeSH)**

Acupuncture Therapy [\*methods]; Antipsychotic Agents [adverse effects] [\*therapeutic use]; Combined Modality Therapy [methods]; Randomized Controlled Trials as Topic; Schizophrenia [\*therapy]

# **MeSH check words**

Humans