

# Review Article

# Acupuncture Therapy for Functional Effects and Quality of Life in COPD Patients: A Systematic Review and Meta-Analysis

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Objective. This study aimed to evaluate the efficacy and safety of acupuncture therapy (AT) for improving functional effects and quality of life in COPD patients. Methods. PubMed, Embase, Cochrane Library, Web of Science, Chinese Biomedical Literature Database (CBM), China National Knowledge Infrastructure (CNKI), Chongqing VIP (CQVIP), and Wanfang Data were searched. The randomized controlled trials (RCTs) evaluating the effect of AT on COPD patients were included. Primary outcome measures included six-minute walk distance (6MWD) and St. George's Respiratory Questionnaire (SGRQ). Study selection, data extraction, and risk of bias assessment were independently conducted, respectively. Statistical analysis was conducted by RevMan software (version 5.3) and Stata software (version 12.0). Results. Nineteen studies (1298 participants) were included. 6MWD improved more (MD: 47.84; 95% CI: 23.33 to 72.35; Z = 3.83, P = 0.0001) and effective rate was higher (OR: 2.26; 95% CI: 1.43 to 3.58; Z = 3.48, P = 0.0005) in the experimental group compared to the control group. Symptom domain scores (MD: -24.86; 95% CI: -32.17 to -17.55; Z = 6.66, P < 0.00001), activity domain scores (MD: -16.52; 95% CI: -22.57 to -10.47; Z = 5.36, P < 0.00001) and impact domain scores (MD: -13.07; 95% CI: -17.23 to -8.92; Z = 6.16, P < 0.00001) of SGRQ in the experimental group improved more compared to the control group. There was no significant improvement in SGRQ total scores between two groups. The improvement of FEV1 was not significant between two groups, yet subgroup analysis showed that patients treated with AT adjunctive to other treatments improved more in FEV<sub>1</sub> (MD: 0.41; 95% CI: 0.28 to 0.54; Z = 6.01, P < 0.00001) compared to those treated with other treatments alone. Conclusion. AT may be effective in improving functional effects and quality of life in COPD patients. Besides, AT may also improve pulmonary function of patients with COPD. However, further high-quality RCTs are needed to confirm the efficacy and safety of AT for COPD patients.

### 1. Introduction

Chronic obstructive pulmonary disease (COPD), a leading cause of morbidity and mortality, is characterized by progressive airflow obstruction, airway inflammation, and systemic effects or comorbidities and is projected to be the third leading cause of death worldwide by 2030 [1, 2]. Since breathlessness, exercise limitation, and health status impairment broadly exist in patients with COPD, effective measures should be taken to improve symptoms, exercise tolerance, and health status based on an individualized assessment of disease [1]. Although appropriate pharmacologic therapy has effect in reducing COPD symptoms and the frequency and severity of exacerbations and improving health status and exercise tolerance [1], its cost and adverse effects can never be ignored.

Acupuncture therapy (AT), one of the most popular treatments in alternative medicine, has been proven to be cost-effective and safe in many conditions [3–5]. However,

there is limited evidence concerning its efficacy and safety. One previous review showed that AT might result in clinically important improvements in quality of life and dyspnea of COPD patients, but it is outdated [6]. Moreover, the interventions of included studies involved point application therapy, acupressure, and transcutaneous electrical stimulation over acupuncture points (Acu-TENS), and these techniques may not genuinely reflect the efficacy of AT based on theories of traditional Chinese medicine. Therefore, the current review aims to evaluate the efficacy and safety of AT for improving functional effects and quality of life in COPD patients.

### 2. Methods

2.1. Inclusion and Exclusion Criteria. We included randomized controlled trials (RCTs) in which the effects of AT on COPD patients were evaluated.

Participants had COPD defined as a clinical diagnosis of COPD, with a postbronchodilator fixed ratio of forced expiratory volume in 1 second (FEV<sub>1</sub>)/forced vital capacity (FVC) < 0.70 measured by spirometry, and those who had an acute exacerbation within four weeks before the study were excluded.

The intervention included AT, such as manual acupuncture, electroacupuncture, auricular acupuncture, and warm acupuncture, yet noninvasive techniques, such as single moxibustion, acupressure, point application, laser acupuncture, or Acu-TENS, were excluded.

Primary outcome measures included any of the following: (i) six-minute walk test/distance (6MWT/6MWD) [26] and (ii) St. George's Respiratory Questionnaire (SGRQ) [27]. Secondary outcome measures included any of the following: (i) FEV<sub>1</sub>, (ii) modified Medical Research Council dyspnea scale (mMRC) [28], (iii) effective rate, and (iv) adverse effects.

2.2. Literature Search. PubMed, Embase, Cochrane Library, Web of Science, Chinese Biomedical Literature Database (CBM), China National Knowledge Infrastructure (CNKI), Chongqing VIP (CQVIP), and Wanfang Data were searched from their inception to 31 December 2017. We developed detailed search strategies for each electronic database without language restrictions. Reference lists of eligible studies and previous systematic reviews were also reviewed to identify further eligible studies.

2.3. Study Selection. Two review authors (Yang Xie and Xueqing Yu) independently examined titles and abstracts retrieved from the search and selected all potentially eligible studies. Then these full-text articles were obtained and the same review authors reviewed them independently against the inclusion and exclusion criteria. A third review author (Jiansheng Li) acted as an arbiter when consensus could not be reached.

2.4. Data Extraction. Data extraction was independently conducted by two review authors (Yang Xie and Xueqing Yu) using a standardized data extraction sheet, involving information of authors, year of publication, study design, participants, intervention, comparator, and outcomes, with a

third review author (Jiansheng Li) acting as an arbiter when disagreements existed between Yang Xie and Xueqing Yu.

2.5. Assessment of Risk of Bias. Methodological quality was evaluated using the Cochrane tool for assessing risk of bias in RCTs [29]. Two review authors (Yang Xie and Xueqing Yu) independently assessed and scored each study with a third review author (Jiansheng Li) acting as an arbiter when disagreements existed.

2.6. Statistical Analysis. Statistical analysis was conducted by RevMan software (version 5.3) [30] and Stata software (version 12.0; StataCorp LP, USA). We summarized data using odds ratio (OR) with 95% confidence intervals (CI) for dichotomous outcomes and mean difference (MD) with 95% CI for continuous outcomes. If the data could not be combined into a meta-analysis, we summarized them in the text. We used a  $\chi^2$  test to estimate heterogeneity of both the MD and OR. Further analysis was performed using the  $I^2$ test. A random-effect model was used to interpret the results if heterogeneity was statistically significant, whereas a fixedeffect model was used if heterogeneity was not statistically significant. We regarded heterogeneity as substantial when  $I^2$  was greater than 50% or a low P value (P < 0.10) was reported for the  $\chi^2$  test [31]. When more than 10 studies were included in the meta-analysis, we would investigate publication bias by funnel plots. In addition, a metaregression analysis was performed to explore potential associations between effect size and covariates of interest (publication year, region, intervention forms, sample size, and treatment period). If necessary, we conducted subgroup analysis to assess whether the treatment effects were different in different subgroups.

### 3. Results

3.1. Literature Search and Study Selection. We retrieved 600 records using the search strategy specified in our protocol. 223 records were discarded after reviewing the titles and/or abstracts. Thirty-five articles that initially appeared to meet the inclusion criteria were excluded with reasons: (i) not stable COPD (n = 24), (ii) not targeted comparators (n = 9), (iii) not targeted outcomes (n = 1), and (iv) full-text articles unavailable (n = 1). Thus, nineteen studies (1298 participants) finally met our criteria and were included in this review [7–25]. The study selection process was outlined in Figure 1.

3.2. Data Extraction and Assessment of Risk of Bias. A detailed description of the characteristics of included studies was outlined in Table 1. We determined the Cochrane "risk of bias" score for each study and this information was summarized in Table 2 and Figures 2 and 3.

#### 3.3. Effects of Interventions

*3.3.1. 6MWD.* Eight studies [9, 10, 16–18, 21, 23, 25] provided numerical data for 6MWD and were included in the meta-analysis. Analysis of the data indicated that there

			TABLE 1: Characteristics of include	d studies.		
Study	Country	Study design	Participants	Interventions	Outcomes	Notes
Chu and Cai [7] 2015	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 15/17, CG: 14/18 Age (range, years): from 46 to 80 FEV <sub>1</sub> % predicted: EG: 56.83 ± 8.96; CG: 52.53 ± 13.96 Course of disease (range): from 5 months to 20 years Participants vere randomly assigned Analyzed: EG: 32 CG: 32	EG: AT + Chinese medicine + western medicine CG: Chinese medicine + western medicine Duration: treatment for 90 days; follow-up for 6 months	Spirometry, acute exacerbation frequency, effective rate, adverse effects	
Deering et al. [8] 2011	Ireland	RCT, 3 arms	Participant status: Gender (M/F): PR group: 11/14; AT + PR group: 8/8; CG: 12/7 Age (years): PR group: $677 \pm 5.3$ ; AT + PR group: $65.1 \pm 9.7$ ; CG: $68.6 \pm 5.5$ FEV <sub>1</sub> % predicted: PR group: $48.5 \pm$ 16.1; AT + PR group: $48.8 \pm 22.7$ ; CG: $45.8 \pm 18.3$ Participants randomly assigned: 60 participants were randomly assigned Analyzed: AT + PR group: 16 PR group: 25	AT + PR group: AT + PR PR group: PR alone CG: no intervention Duration: treatment for 7 weeks; follow-up for 3 months	Body mass index, mMRC, the modified Borg dyspnea score, systemic inflammation, spirometry, total energy expenditure, physical activity duration, metabolic equivalents, steps per day, sleep time/efficiency, incremental shuttle walk test, SGRQ, EQ-5D	Control group was not used in the analysis
Deng et al. [9] 2016	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 13/9, CG: 15/7 Age (years): EG: 57.1 $\pm$ 5.9; CG: 58.5 $\pm$ 6.5 FEV <sub>1</sub> % predicted: EG: 52.4 $\pm$ 2.9; CG: 51.5 $\pm$ 2.7 Course of disease (years): EG: 6.4 $\pm$ 2.5; CG: 6.3 $\pm$ 2.1 Participants vere randomly assigned Analyzed: EG: 22 CG: 22	EG: abdominal AT + conventional therapy CG: conventional therapy alone Duration: treatment for 24 weeks	Annual hospital stay, annual acute exacerbation frequency, oxygen saturation level, spirometry, 6MWD	

			TABLE 1: Continued.			
Study	Country	Study design	Participants	Interventions	Outcomes	Notes
Feng et al. [10] 2016	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 33/3, CG: 31/5 Age (years): EG: 67.8 $\pm$ 5.4; CG: 67.1 $\pm$ 6.1 FEV <sub>1</sub> % predicted: EG: 47.3 $\pm$ 17.1; CG: 43.9 $\pm$ 16.8 Participants randomly assigned: 72 participants were randomly assigned Analyzed: EG: 33 CG: 32	EG: AT + daily medication CG: Sham acupuncture + daily medication Duration: treatment 3 times weekly for 8 weeks	6MWD, modified Borg scale score before and after 6MWT, oxygen saturation during the 6MWT; spirometry, SGRQ	
Gao et al. [11] 2011	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 13/17, CG: 12/18 Age (years): EG: 64.87 $\pm$ 8.73; CG: 65.25 $\pm$ 10.66 FEV <sub>1</sub> % predicted: EG: 45.88 $\pm$ 5.05; CG: 43.54 $\pm$ 6.29 Course of disease (years): EG: 10.31 $\pm$ 5.82; CG: 10.78 $\pm$ 5.53 Participants randomly assigned: 61 participants were randomly assigned Analyzed: EG: 30 CG: 30	EG: warm AT CG: drug therapy Duration: treatment for 8 weeks	Spirometry, clinical symptoms, SGRQ	
Hu [12] 2016	China	RCT, 2 arms	Participant status: Gender (M/F): not described Age (years): not described FEV <sub>1</sub> % predicted: not described Course of disease (years): not described Participants randomly assigned: 89 participants were randomly assigned Analyzed: EG: 45 CG: 44	EG: warm AT + conventional therapy CG: spreading moxibustion + conventional therapy Duration: treatment for 2 months	Effective rate, spirometry	

Outcomes Notes	+ rugs Trugs AT + PR Ffective rate, spirometry AT + PR ffective rate, spirometry drugs ment for ment for PR group were analysis	+ drug Safety indicators, CAT, spirometry, clinical symptoms, effective rate, ment for 1 adverse effects	ese stern Clinical symptoms, CAT, edicine + spirometry, effective
Interventions	AT group: AT + conventional dru PR group: PR + conventional dru AT + PR group: / + conventional d Duration: treatm 100 days	EG: warm AT + 6 therapy CG: drug therapy Duration: treatm month	EG: AT + Chines medicine + west medicine CG: Chinese medicin western medicin
TABLE 1: Continued. Particinants	Participants status: Participant status: Gender (M/F): AT group: 9/13; PR group: 8/14; AT + PR group: 10/12 Age (years): AT group: 10/12 Age (years): AT group: 10/12 PR group: 60.0 $\pm$ 34.8; AT + PR group: 61.0 $\pm$ 33.2 FEV <sub>1</sub> % predicted: AT group: 70.9 $\pm$ 0.2; PR group: 71.1 $\pm$ 0.1; AT + PR group: 70.6 $\pm$ 0.7 Course of disease (years): AT group: 11.7 $\pm$ 8.9; PR group: 12.1 $\pm$ 7.2; AT + PR group: 11.6 $\pm$ 9.0 Participants randomly assigned: 66 participants were randomly assigned Analyzed: AT PR group: 22 AT + PR group: 22	Participant status: Gender (M/F): EG: 19/10, CG: 17/13 Age (years): EG: 57.21 $\pm$ 6.68; CG: 55.80 $\pm$ 7.23 FEV <sub>1</sub> % predicted: EG: 66.28 $\pm$ 6.86; CG: 65.16 $\pm$ 6.16 Course of disease (years): EG: 10.38 $\pm$ 4.90; CG: 10.70 $\pm$ 4.88 Participants randomly assigned: 60 participants were randomly assigned Analyzed: EG: 29 CG: 30	<i>Participant status:</i> Gender (M/F): two groups were reported to be comparable in gender. Age (years): two groups were reported to be comparable in age. FEV <sub>1</sub> % predicted: EG: 53.1 ± 10.9; CG: 51.9 ± 11.4 <i>Participants randomly assigned:</i>
Study design	RCT, 3 arms	RCT, 2 arms	RCT, 2 arms
Country	China	China	China
Study	Jia [13] 2004	Li [14] 2015	Li et al. [15] 2016

StudyCountryStudy designParticipantsOutcomesNiList et al.CountryStudy designParticipantsOutcomesNiList et al.ChinaRC(3, 84, 15, 65, 35/1 ± 23, 65, 53/1 ± 24, 66, 66, 26/14EG, AT 4 rug therapyCinical signs and2015CG: 36, 45, 46, 56, 53/1 ± 23, 75, 52, 53/1 ± 24, 66, 56/14Duration: treinieri forspinonetry, effective rate2015CG: 36, 45, 46, 50, 50, 54/1Duration: treinieri forspinonetry, effective rate2015CG: 36, 45, 46, 52, 312, 45, 56, 55, 512, 52, 56, 56, 52, 52, 54Duration: treinieri forspinonetry, effective rate2012AnalysisDuration: treinieri forspinonetry, effective ratesale core2012Ci 40Ci 40Ci 53, 512, 52, 55, 512, 52, 55, 512, 52, 52, 55Ci 74, 4 daiySON, Ti spinonetry, effective rate2012JapanRCI, 2 amsRCI, 2 amsRCI, 2 amsSouth rateSouth rate2012JapanRCI, 2 amsRCI, 2 amsRCI, 2 amsSouth rate2012JapanRCI, 2 amsRCI, 2 amsSouth rateSouth rate2012JapanRCI, 2 amsRCI, 2 amsSouth rateSouth rate2012JapanRCI, 2 amsRCI, 2 amsRCI, 2 amsSouth rate2013JapanRCI, 2 amsRCI, 2 amsSouth rateSouth rate2014RCI, 2 amsRCI, 2 amsRCI, 2 amsSouth rateSouth rate2015JapanRCI, 2 amsRCI, 2 amsSouth rateSouth ra				TABLE 1: Continued.			
List et al. [6]ChinaRCT.2 ams Rg (varo): EC: 33 ± 12.4 CG: 652.4 G (Sol ± 6.42) FFV % predictorh EC: 35.1 ± 7.28, G (Sol ± 6.42) FFV % predictorh EC: 35.1 ± 7.28, FEV % predictorh South 9.3 stepticipants were randomly signed.EG: AT + drug theopy for signed. 3 monthsGinal signs and syntoms, SOVD, modified Borg evaluation2015ChinaRCT.2 ams Supercipants were randomly signed.Stepticipants were randomly signed.EG: AT + drug theopy for syntoms, SOVD, modified BorgStepticipants were randomly signed.2012Spanki et al. [7]JapanRCT.2 ams Stepticipants were randomly signed.Stepticipants were randomly signed.Stepticipants were randomly signed.2012JapanRCT.2 ams Stepticipants were randomly signed.RCMT + drug theopy for signed.Stepticipants were randomly signed.Stepticipants signed.2012JapanRCT.2 amsRCT.2 ams Stepticipants were randomly signed.Stepticipants were randomly signed.Stepticipants signed.Stepticipants signed.2012JapanRCT.2 amsRCT.2 amsRCT.2 ams signed.Stepticipants signed.Stepticipants signed.Stepticipants signed.Stepticipants signed.2012JapanRCT.2 amsRCT.2 ams signed.RCT.2 ams signed.Stepticipants signed.Stepticipants signed.Stepticipants signed.2013JapanRCT.2 amsRCT.2 ams signed.RCT.2 ams signed. <th>Study</th> <th>Country</th> <th>Study design</th> <th>Participants</th> <th>Interventions</th> <th>Outcomes</th> <th>Notes</th>	Study	Country	Study design	Participants	Interventions	Outcomes	Notes
Brucki et al. [17] Japan RCT, 2 arms Participant states: Gender (MIP): EG: 31/3 CG: 32/2 Age (years): EG: 72.5 ± EG: AT + daily Age (years): EG: 61 ± 10.2 ± EG: AT + aterolic EG: AT + aterolic EG: AT + aterolic   Tong et al. [8] China RCT, 2 arms Participant states EG: AT + aterolic EG: AT + aterolic EG: AT + aterolic   Tong et al. [8] China RCT, 2 arms Participant states EG: AT + aterolic EG: AT + aterolic EG: AT + aterolic   Tong et al. [8] China RCT, 2 arms Participants were randomly EG: AT + aterolic EG: AT + aterolic EG: AT + aterolic   Tong et al. [8] China RCT, 2 arms Participants were randomly EG:	Liu et al. [16] 2015	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 24/16, CG: 26/14 Age (years): EG: 58.3 $\pm$ 12.4; CG: 63.2 $\pm$ 10.7 FEV <sub>1</sub> % predicted: EG: 35.71 $\pm$ 7.28; CG: 36.42 $\pm$ 6.42 Participants randomly assigned: 80 participants were randomly assigned Analyzed: EG: 40 CG: 40	EG: AT + drug therapy CG: drug therapy alone Duration: treatment for 3 months	Clinical signs and symptoms, 6MWD, spirometry, effective rate	
Tong et al. [18] Tong et al. [19] Tong et al. [10] Tong et al. [10] Tong et al. [18] Tong et al. [19] Tong et al. [18] Tong et al. [19] Tong et al. [18] Tong et al. [19] Tong et al. [10] Tong et al. [11] Ton	Suzuki et al. [17] 2012	Japan	RCT, 2 arms	Participant status: Gender (M/F): EG: $31/3$ , CG: $32/2$ Age (years): EG: $72.7 \pm 6.8$ ; CG: $72.5 \pm 7.4$ FEV <sub>1</sub> % predicted: EG: $44.5 \pm 16.3$ ; CG: $48.0 \pm 16.5$ Participants randomly assigned: 68 participants were randomly assigned Analyzed: EG: 30 CG: 32	EG: AT + daily medication CG: placebo acupuncture + daily medication Duration: treatment once a week for 12 weeks	6MWD, modified Borg scale score before and after 6MWT, oxygen saturation during the 6MWT, spirometry, SGRQ, arterial blood gas, maximum inspiratory mouth pressure, maximum expiratory mouth pressure, range of motion in the rib cage, body mass index, serum prealbumin levels, MRC score, adverse reactions	
	Tong et al. [18] 2014	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 15/1, CG: 12/2 Age (years): EG: $64 \pm 6;$ CG: $67 \pm 6$ FEV <sub>1</sub> % predicted: EG: $41.72 \pm 17.95;$ CG: $36.16 \pm 16.29$ Participants randomly assigned: 30 participants were randomly assigned Analyzed: EG: 16 CG: 14	EG: AT + aerobic exercise CG: placebo acupuncture + aerobic exercise Duration: treatment for 5 weeks	6MWD, spirometry, maximum oxygen uptake, exercise time, SGRQ	

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udy Country Study design Participants In   and tet al. [19] Country Study design Participants attats: 12.77;   and tet al. [19] China RCT, 2 arms Gender (MJP): EG: 47/28, CG: 45/30 20   and tet al. [19] China RCT, 2 arms Son CG: 47.27 ± 14.02 20   0.9 a.7.27 ± 14.02 EG: 75 EG 20   0.9 CG: 47.27 ± 14.02 EG: 75 20   0.0 CG: 47.27 ± 14.02 EG: 75 20   1.0 Dariticipants were randomly assigned. 36   2.0 Son CG: 779 ± 29.0 20   2.0 Age (years): EG: 18/3 ± 5.06 EG   0.14 China RCT, 2 arms 20   0.15 China RCT, 2 arms 20   10.1 China RCT, 2 arms 20   10.2 China RCT, 2 arms 20   10.3 EG: 40 CHINPERC: 18/3 ± 5.06   10.4 China <		TABLE 1: Continued.			
net right status: Participant status:   Gender (M/F): EG: 47/8, CG: 45/30   Age (years): EG: 62.40 ± 8.56; CG:   0.0   0.1   1.1   1.1   1.2   1.3   1.4   1.4   1.5   1.5   1.6   1.7   1.7   1.8   1.9   1.9   1.1   1.1   1.2   1.2   1	Country Study des	ign Participants	Interventions	Outcomes	Notes
Participant status: Participant status:   Gender (MrF): EG: 22/18, CG: 18/22   Age (years): EG: 68.9 ± 8.7; CG: 68.5 ± 9, 68   Gender (MrF): EG: 22/18, CG: 18/22   Gender (MrF): EG: 22/18, CG: 68.5 ± 9, 506;   CG: 43.55 ± 6.30   CG: 43.55 ± 6.30   Course of disease (years): EG: 11.8 ± 0, 0   Dipparticipants randomly assigned:   81   B2   Participants randomly assigned:   B3   B4   Analyzed:   EG: 40   CG: 40   CG: 40   CG: 40   Darticipant status:   Gender (M/F): EG: 26/14, CG: 28/12   Age (range, years): EG: from 49 to 78;   CG: from 45 to 77   B4/16   B5   CG: from 45 to 77   FEV16   CO   CG: from 45 to 77   FEV16   CG: from 45 to 77   B1   CD   CG: from 45 to 77   FEV16   S0 participants randomly assigned:   Age (range, years): EG: 60, CG: 21   D1   CD   CG: from 45 to 77 <td< td=""><td>China RCT, 2 ar</td><td>Participant status: Gender (M/F): EG: <math>47/28</math>, CG: <math>45/30</math> Age (years): EG: <math>62.40 \pm 8.56</math>; CG: <math>61.80 \pm 10.10</math> FEV<sub>1</sub>% predicted: EG: <math>48.07 \pm 12.77</math>; CG: <math>47.27 \pm 14.02</math> Course of disease (years): EG: <math>7.73 \pm 3.80</math>; CG: <math>7.70 \pm 2.92</math> Participants randomly assigned: 150 participants were randomly assigned Analyzed: EG: 75 CG: 75 CG: 75</td><td>EG: AT + Chinese medicine CG: Chinese medicine alone Duration: treatment for 36 days</td><td>Effective rate, spirometry</td><td></td></td<>	China RCT, 2 ar	Participant status: Gender (M/F): EG: $47/28$ , CG: $45/30$ Age (years): EG: $62.40 \pm 8.56$ ; CG: $61.80 \pm 10.10$ FEV <sub>1</sub> % predicted: EG: $48.07 \pm 12.77$ ; CG: $47.27 \pm 14.02$ Course of disease (years): EG: $7.73 \pm 3.80$ ; CG: $7.70 \pm 2.92$ Participants randomly assigned: 150 participants were randomly assigned Analyzed: EG: 75 CG: 75 CG: 75	EG: AT + Chinese medicine CG: Chinese medicine alone Duration: treatment for 36 days	Effective rate, spirometry	
$\begin{array}{llllllllllllllllllllllllllllllllllll$	China RCT, 2 ar	Participant status:Gender (M/F): EG: 22/18, CG: 18/22Age (years): EG: 68.9 $\pm$ 8.7; CG: 68.5 $\pm$ 9.6FEV1% predicted: EG: 45.89 $\pm$ 5.06;CG: 43.55 $\pm$ 6.30Course of disease (years): EG: 11.8 $\pm$ 6.5; CG: 12.3 $\pm$ 5.5Participants randomly assigned:80 participants were randomlyassignedAnalyzed:EG: 40CG: 40CG: 40	EG: warm AT CG: drug therapy Duration: treatment for 8 weeks	Spirometry, symptom scores	
assigned Analyzed: EG: 40 CG: 40	China RCT, 2 ar	Participant status:Gender (M/F): EG: 26/14, CG: 28/12Age (range, years): EG: from 49 to 78;CG: from 45 to 77EEV1% predicted: EG: 53.5 $\pm$ 19.2; CG:54.3 $\pm$ 22.6Course of disease (mean, years): EG:6.9; CG: 71Participants vere randomlyassignedAnalyzed:EG: 40CG: 40CG: 40	EG: AT + PR CG: PR alone Duration: treatment for 40 days	COPD quality of life questionnaire, 6MWD, spirometry	

			TABLE 1: Continued.			
Study	Country	Study design	Participants	Interventions	Outcomes	Notes
Yu [22] 2014	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 18/12, CG: 17/13 Age (years): EG: 63.0 $\pm$ 8.5; CG: 62.0 $\pm$ 7.6 FEV <sub>1</sub> % predicted: EG: 50.23 $\pm$ 2.56; CG: 51.33 $\pm$ 2.43 Course of disease (years): EG: 8.9 $\pm$ 3.7; CG: 8.4 $\pm$ 3.5 Participants randomly assigned: 60 participants were randomly assigned Analyzed: EG: 30 CG: 30	EG: warm AT + function training CG: conventional drug therapy + function training Duration: treatment for 3 months	Arterial blood gas, spirometry, SGRQ, effective rate	
Ge et al. [23] 2017	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 23/1, CG: 15/5 Age (years): EG: $65 \pm 6$ ; CG: $65 \pm 7$ FEV <sub>1</sub> % predicted: EG: $40.76 \pm 16.36$ ; CG: $40.53 \pm 17.40$ Course of disease (years): EG: $9.1 \pm 5.5$ ; CG: $8.6 \pm 6.8$ Participants vandomly assigned: 44 participants were randomly assigned Analyzed: EG: 22 CG: 19 CG: 19	EG: AT + conventional drugs + aerobic exercise CG: placebo acupuncture + conventional drugs + aerobic exercise Duration: treatment for 14 times	Body mass index, average distance and average maximum heart rate during bicycle exercise, 6MWD, maximum power and maximum heart rate during exercise cardiopulmonary function test, spirometry	

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			TABLE 1: Continued.			
Study	Country	Study design	Participants	Interventions	Outcomes	Notes
Shi [24] 2017	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 18/12, CG: 17/14 Age (years): EG: 57.77 $\pm$ 6.54; CG: 55.90 $\pm$ 6.86 FEV <sub>1</sub> % predicted: EG: 64.11 $\pm$ 5.79; CG: 65.85 $\pm$ 6.86 Course of disease (years): EG: 11.43 $\pm$ 4.37; CG: 11.68 $\pm$ 3.64 Participants randomly assigned: 66 participants were randomly assigned Analyzed: EG: 30 CG: 31	EG: AT CG: western medicine Duration: treatment for 2 months	CAT, spirometry, clinical symptoms, effective rate, safety indicators	
Tang [25] 2017	China	RCT, 2 arms	Participant status: Gender (M/F): EG: 19/11, CG: 18/13 Age (years): EG: 57.47 $\pm$ 5.33; CG: 56.59 $\pm$ 6.34 Course of disease (years): EG: 10.07 $\pm$ 2.20; CG: 10.55 $\pm$ 2.10 Participants were randomly assigned: Analyzed: EG: 30 CG: 29	EG: AT + western medicine CG: western medicine alone Duration: treatment for 2 months	CAT, clinical symptoms, 6MWT, effective rate, safety indicators	
RCT: randomized contro test/distance, SGRQ: St. C	lled trial, EG: experi George's Respiratory	mental group, CG: contrc Questionnaire, mMRC: 1	ol group, AT: acupuncture therapy, PR: pulmonary modified Medical Research Council dyspnea scale	rehabilitation, FEV <sub>1</sub> : forced expira , MRC: Medical Research Council	tory volume in 1 second, 6MWT/MWD dvspnea scale. CAT: COPD assessment	D: six-minute walk

Study	Random sequence generation	Allocation concealment	Blinding of participants and personnel	Blinding of outcome assessment	Incomplete outcome data	Selective reporting	Other biases
Chu and Cai [7] 2015	U	U	Н	U	L	U	Н
Deering et al. [8] 2011	L	U	Н	L	U	U	Н
Deng et al. [9] 2016	U	U	Н	U	L	U	Н
Feng et al. [10] 2016	L	L	L	L	L	U	L
Gao et al. [11] 2011	L	U	Н	U	L	U	L
Hu [12] 2016	U	U	Н	U	L	U	L
Jia [13] 2004	L	U	Н	U	L	U	L
Li [14] 2015	L	U	Н	U	L	U	Н
Li et al. [15] 2016	L	L	Н	U	L	U	L
Liu et al. [16] 2015	U	U	Н	U	L	U	Н
Suzuki et al. [17] 2012	L	U	L	L	L	U	L
Tong et al. [18] 2014	L	L	L	L	L	U	L
Wan et al. [19] 2009	U	U	Н	U	L	U	L
Xie and Yu [20] 2014	L	U	Н	U	L	U	L
Yang et al. [21] 2009	Н	U	Н	U	L	U	Н
Yu [22] 2014	L	U	Н	U	L	Н	L
Ge et al. [23] 2017	L	U	L	U	L	U	L
Shi [24] 2017	L	U	Н	U	L	U	Н
Tang [25] 2017	L	U	Н	U	L	U	Н

TABLE 2: Risks of bias of included studies.

Notes. Quality assessment based on the Cochrane tools for assessing risk of bias. L: low (low risk of bias), H: high (high risk of bias), U: unclear (uncertain risk of bias).

was heterogeneity ( $\chi^2 = 65.96$ , P < 0.00001;  $I^2 = 89\%$ ); hence, a random-effect model was used. The pooled results showed that 6MWD in the experimental group improved more compared to the control group (MD: 47.84; 95% CI: 23.33 to 72.35; Z = 3.83, P = 0.0001) (Figure 4).

3.3.2. SGRQ. Five studies [8, 10, 11, 17, 18] provided numerical data for SGRQ total scores and were included in the metaanalysis. Analysis of the data indicated that there was heterogeneity ( $\chi^2 = 39.18$ , P < 0.00001;  $I^2 = 90\%$ ); hence, a random-effect model was used. The pooled results showed that there was no significant improvement in SGRQ total scores between two groups (MD: -6.58; 95% CI: -13.19 to 0.03; Z = 1.95, P = 0.05) (Figure 5).

Two studies [10, 17] provided numerical data for symptom domain scores of SGRQ and were included in the metaanalysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 0.05$ , P = 0.83;  $I^2 = 0\%$ ); hence, a fixed-effect model was used. The pooled results showed that symptom domain scores of SGRQ in the experimental group improved more compared to the control group (MD: -24.86; 95% CI: -32.17 to -17.55; Z = 6.66, P < 0.00001) (Figure 6).

Two studies [10, 17] provided numerical data for activity domain scores of SGRQ and were included in the metaanalysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 0.04$ , P = 0.84;  $I^2 = 0\%$ ); hence, a fixed-effect model was used. The pooled results showed that activity domain scores of SGRQ in the experimental group improved more compared to the control group (MD: -16.52; 95% CI: -22.57 to -10.47; Z = 5.36, P < 0.00001) (Figure 7).

Two studies [10, 17] provided numerical data for impact domain scores of SGRQ and were included in the metaanalysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 0.00$ , P = 0.97;  $I^2 = 0\%$ );



FIGURE 1: Study flow diagram.

hence, a fixed-effect model was used. The pooled results showed that impact domain scores of SGRQ in the experimental group improved more compared to the control group (MD: -13.07; 95% CI: -17.23 to -8.92; Z = 6.16, P < 0.00001) (Figure 8).

3.3.3. *FEV*<sub>1</sub>. Seven studies [7, 11, 12, 17, 19, 20, 22] provided numerical data for FEV<sub>1</sub> and were included in the metaanalysis. Analysis of the data indicated that there was heterogeneity ( $\chi^2 = 30.40$ , P < 0.0001;  $I^2 = 80\%$ ); hence, a random-effect model was used. The pooled results showed that there was no significant improvement in FEV<sub>1</sub> between two groups (MD: 0.13; 95% CI: -0.05 to 0.31; Z = 1.44, P =0.15) (Figure 9).

3.3.4. *Effective Rate.* Ten studies [7, 12–16, 19, 22, 24, 25] provided categorical data for effective rate and were included

in the meta-analysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 8.33$ , P = 0.50;  $I^2 = 0\%$ ); hence, a fixed-effect model was used. The pooled results showed that effective rate in the experimental group was higher compared to the control group (OR: 2.26; 95% CI: 1.43 to 3.58; Z = 3.48, P = 0.0005) (Figure 10).

3.3.5. *mMRC*. Only one study [8] provided numerical data for mMRC scores; thus, the meta-analysis was not performed. Changes from baseline in mMRC scores in AT plus pulmonary rehabilitation (PR) group and PR group were  $-0.3 \pm 0.5$  and  $-0.3 \pm 0.9$ , respectively. There was significant difference reported within AT plus PR group (P = 0.04).

*3.4. Adverse Effects.* Six studies [7, 14, 15, 17, 24, 25] provided information about adverse effects. Only one study [17] reported some minor adverse reactions during the trial



FIGURE 2: Risk of bias summary: review authors' judgements about each risk of bias item for each included study.

including fatigue, subcutaneous hemorrhage, dizziness, and needle site pain, and the remaining 5 studies [7, 14, 15, 24, 25] reported no adverse effects.

3.5. *Metaregression Analysis*. We tried to perform a univariate metaregression analysis to explore potential associations

between effect size and covariates of interest (publication year, region, intervention forms, sample size, and treatment period) (see Table 3). However, the results showed that there were no statistically significant associations among them, and this might be due to the insufficient number of studies included [32].



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

Cturden an Carl announ	Ex	perimer	ntal	(	Control	l	XA7. :	Mean Difference		Mean	Difference		
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI		IV, Ran	dom, 95% C	I	
Deng et al. 2016	49.4	42.26	22	20.1	46.17	22	12.3%	29.30 [3.15, 55.45]					
Feng et al. 2016	67.4	21.7	33	-13.8	54.8	32	13.0%	81.20 [60.82, 101.58]					
Ge et al. 2017	58.05	51.52	22	19.03	48.18	19	11.7%	39.02 [8.48, 69.56]					
Liu et al. 2015	53.1	34.27	40	44.7	30.81	40	13.7%	8.40 [-5.88, 22.68]			+ <b>-</b> -		
Suzuki et al. 2012	63.5	49.9	30	-19.4	48.7	32	12.5%	82.90 [58.33, 107.47]			-		
Tang 2017	119.13	42.26	30	34.07	46.17	29	12.8%	85.06 [62.45, 107.67]			-		
Tong et al. 2014	49.23	53.9	16	11.87	48.35	14	10.9%	37.36 [0.77, 73.95]				-	
Yang et al. 2009	43.1	42.26	40	23	46.17	40	13.1%	20.10 [0.70, 39.50]					
Total (95% CI)			233			228	100.0%	47.84 [23.33, 72.35]					
Heterogeneity: $\tau^2 =$	1091.89;	$\chi^2 = 6$	5.96, df	= 7 (P	< 0.00	001); I	$^{2} = 89\%$		-200	-100	0	100	200
Test for overall effect	:: Z = 3.8	83 ( <i>P</i> =	0.0001	)						Favours [control]	Favours	[experime	ntal]

FIGURE 4: Experimental group versus control group, 6MWD.

Starlar an Salamann	Exp	erimer	ntal	(	Contro	1	<b>XAZ-:</b> -1-4	Mean Difference		Me	an Diffe	rence	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI		IV, R	andom,	95% CI	
Deering et al. 2011	-7.1	12.7	16	-7.4	8.7	25	18.9%	0.30 [-6.80, 7.40]				_	
Feng et al. 2016	-15.3	24.8	33	0.5	20	32	14.5%	-15.80 [-26.74, -4.86]			-		
Gao et al. 2011	-12.84	3.43	30	-10.21	3.89	30	23.6%	-2.63 [-4.49, -0.77]			-		
Suzuki et al. 2012	-16	9.7	30	0.3	7.9	32	21.7%	-16.30 [-20.72, -11.88]					
Tong et al. 2014	-4.13	4.77	16	-3.29	8.28	14	21.2%	-0.84[-5.77, 4.09]			-		
Total (95% CI)			125			133	100.0%	-6.58 [-13.19, 0.03]		•			
Heterogeneity: $\tau^2 =$	47.16; χ	$^{2} = 39$	.18, df =	= 4 (P <	0.000	01); $I^2$	= 90%	-	-50	-25	0	25	50
Test for overall effec	t: $Z = 1.9$	95 (P =	= 0.05)						Favo	urs [experimer	ntal]	Favours [control]	

FIGURE 5: Experimental group versus control group, SGRQ total scores.

Star las en Sals annos	Exp	erime	ntal	(	Contro	ol	147. :l. 4	Mean Difference		Me	an Dif	ference	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Fixed, 95% CI		IV,	Fixed,	95% CI	
Feng et al. 2016	-25.4	44.8	33	-2.4	30.8	32	15.4%	-23.00 [-41.64, -4.36]	_	-	_		
Suzuki et al. 2012	-27.8	17.3	30	-2.6	14.4	32	84.6%	-25.20 [-33.15, -17.25]					
Total (95% CI)			63			64	100.0%	-24.86 [-32.17, -17.55]		$\bullet$			
Heterogeneity: $\chi^2$ = Test for overall effec	0.05, df t: <i>Z</i> = 6.	= 1 (. 66 (P	P = 0.83 < 0.000	3); $I^2 = 001$ )	0%			-	-50 Favou	–25 Irs [experimen	0 ntal]	25 Favours [cont	+ 50 rrol]

FIGURE 6: Experimental group versus control group, symptom domain scores of SGRQ.

Study or Subgroup	Exp	perime	ntal	(	Contro	ol	Weight	Mean Difference		Mean	Difference		
Study of Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Fixed, 95% CI		IV, Fix	ed, 95% CI		
Feng et al. 2016 Suzuki et al. 2012	-13.7 -14	24.1 13.6	33 30	1.5 2.8	34.4 13.1	32 32	17.4% 82.6%	-15.20 [-29.68, -0.72] -16.80 [-23.45, -10.15]		-	—		
Total (95% CI)			63			64	-16.52 [-22.57, -10.47]		-				
Heterogeneity: $\chi^2$ = Test for overall effect	0.04, df t: Z = 5.	f = 1 (1) .36 (P)	P = 0.8 < 0.000	4); $I^2 =$ 001)	0%				-50 Favor	-25 1rs [experimental	0 [] Favou	25 rs [control]	50



Study or Subgroup	Exp	erime	ntal	Control			Maight	Mean Difference		Mean	Differ	rence	
	Mean	SD	Total	Mean	SD	Total	weight	IV, Fixed, 95% CI		IV, Fix	ked, 95	5% CI	
Feng et al. 2016	-12.5	25.4	33	0.4	21.9	32	13.0%	-12.90 [-24.42, -1.38]			-		
Suzuki et al. 2012	-13	9.6	30	0.1	8.2	32	87.0%	-13.10 [-17.56, -8.64]					
Total (95% CI)			63			64	100.0%	-13.07 [-17.23, -8.92]		•			
Heterogeneity: $\chi^2 = 0.00$ , df = 1 ( <i>P</i> = 0.97); $I^2 = 0\%$										-25	0	25	50
Test for overall effect: $Z = 6.16$ ( $P < 0.00001$ )									Favo	ours [experimental	l]	Favours [control]	

FIGURE 8: Experimental group versus control group, impact domain scores of SGRQ.

Study on Sub moun	Exp	perimei	ntal		Control		Mainht	Mean Difference		Mean	Differen	ice	
Study of Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI		IV, Rano	% CI		
Chu and Cai 2015	0.53	0.43	32	0.12	0.56	32	13.8%	0.41 [0.17, 0.65]			-		
Gao et al. 2011	0.23	0.56	30	0.39	0.92	30	9.9%	-0.16 [-0.55, 0.23]					
Hu 2016	0.07	0.43	45	0.14	0.56	44	14.9%	-0.07 [-0.28, 0.14]					
Suzuki et al. 2012	0.07	0.3	30	-0.04	0.2	32	17.1%	0.11 [-0.02, 0.24]					
Wan et al. 2009	0.51	0.43	75	0.1	0.56	75	16.2%	0.41 [0.25, 0.57]					
Xie and Yu 2014	0.23	0.43	40	0.39	0.56	40	14.6%	-0.16 [-0.38, 0.06]			+		
Yu 2014	0.1	0.43	30	-0.18	0.56	30	13.5%	0.28 [0.03, 0.53]					
Total (95% CI)			282			283	100.0%	0.13 [-0.05, 0.31]					
Heterogeneity: $\tau^2 = 0.04$ ; $\chi^2 = 30.40$ , df = 6 (P < 0.0001); I <sup>2</sup> = 80%										-0.5	0	0.5	1
Test for overall effect: $Z = 1.44$ ( $P = 0.15$ )										Favours [control]	Favo	urs [experime	ntal]

FIGURE 9: Experimental group versus control group,  $FEV_1$ .

	Experin	nental	Con	trol		Odds Ratio		Odd	ls Ratio		_		
Study or Subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% C	Ι	M-H, Fiz	M-H, Fixed, 95% CI				
Chu and Cai 2015	30	32	24	32	6.0%	5.00 [0.97, 25.77]			-				
Hu 2016	39	45	41	44	22.0%	0.48 [0.11, 2.03]			<u> </u>				
Jia 2004	41	44	17	22	6.2%	4.02 [0.86, 18.73]							
Li 2015	26	29	25	30	10.1%	1.73 [0.37, 8.03]							
Li et al. 2016	28	30	27	30	7.2%	1.56 [0.24, 10.05]							
Liu et al. 2015	38	40	32	40	6.4%	4.75 [0.94, 23.98]							
Shi 2017	26	30	25	31	13.1%	1.56 [0.39, 6.20]							
Tang 2017	26	30	22	29	11.9%	2.07 [0.53, 8.00]							
Wan et al. 2009	71	75	66	75	14.0%	2.42 [0.71, 8.24]		-					
Yu 2014	29	30	24	30	3.2%	7.25 [0.82, 64.46]							
Total (95% CI)		385		363	100.0%	2.26 [1.43, 3.58]			•				
Total events	354		303						-	L			
Heterogeneity: $\chi^2 = 3$	8.33, df =	9 $(P = 0.5)$	50); $I^2 = 0$	%			0.01	0.1	1 1	0 100			
Test for overall effect: $Z = 3.48$ ( $P = 0.0005$ )								Favours [control]	Favours [ex	perimental]			

FIGURE 10: Experimental group versus control group, effective rate.

Committee of interest	Coefficient	Standard amon (SE)	4	ת	95% confider	nce intervals (CI)
Covariates of interest	Coemcient	Standard error (SE)	t	P	Lower limit	Upper limit
Six-minute walk distance						
Publication year	2.812	4.479	0.63	0.553	-8.148	13.773
Region	*	*	*	*	*	*
Intervention forms	26.947	21.520	1.25	0.257	-25.710	79.603
Sample size	-0.236	0.716	-0.33	0.753	-1.988	1.516
Treatment period	-0.115	0.292	-0.39	0.707	-0.829	0.599
SGRQ						
Publication year	-1.634	2.107	-0.78	0.495	-8.340	5.072
Region	8.595	9.520	0.90	0.433	-21.702	38.891
Intervention forms	4.605	4.389	1.05	0.371	-9.363	18.573
Sample size	-0.382	0.225	-1.70	0.188	-1.097	0.334
Treatment period	-0.342	0.139	-2.47	0.090	-0.783	0.099
$FEV_1$						
Publication year	-0.028	0.043	-0.65	0.544	-0.139	0.083
Region	*	*	*	*	*	*
Intervention forms	-0.086	0.088	-0.98	0.371	-0.313	0.140
Sample size	0.003	0.003	0.83	0.447	-0.005	0.011
Treatment period	0.002	0.005	0.48	0.654	-0.011	0.016
Effective rate						
Publication year	-0.064	0.060	-1.07	0.315	-0.203	0.074
Region	*	*	*	*	*	*
Intervention forms	-0.117	0.301	-0.39	0.707	-0.812	0.577
Sample size	-0.002	0.008	-0.22	0.830	-0.021	0.017
Treatment period	0.015	0.010	1.54	0.163	-0.008	0.037

TABLE 3: Univariate metaregression analysis of covariates of interest.

Note. 6MWD: six-minute walk test/distance, SGRQ: St. George's Respiratory Questionnaire. \* The region was the same.

# 3.6. Subgroup Analysis (AT Adjunctive to Other Treatments versus Placebo or Sham Acupuncture Adjunctive to Other Treatments)

3.6.1. 6MWD. Four studies [10, 17, 18, 23] provided numerical data for 6MWD and were included in the meta-analysis. Analysis of the data indicated that there was heterogeneity ( $\chi^2 = 9.17$ , P = 0.03;  $I^2 = 67\%$ ); hence, a random-effect model was used. The pooled results showed that 6MWD in the experimental group improved more compared to the control group (MD: 63.05; 95% CI: 39.27 to 86.83; Z = 5.20, P < 0.00001) (Figure 11).

3.6.2. SGRQ. Three studies [10, 17, 18] provided numerical data for SGRQ total scores and were included in the metaanalysis. Analysis of the data indicated that there was heterogeneity ( $\chi^2 = 22.16$ , P < 0.0001;  $I^2 = 91\%$ ); hence, a random-effect model was used. The pooled results showed that SGRQ total scores in the experimental group improved more compared to the control group (MD: -10.66; 95% CI: -22.24 to 0.92; Z = 1.80, P = 0.07) (Figure 12).

3.6.3.  $FEV_1$ . Only one study [17] provided numerical data for  $FEV_1$ ; thus, the meta-analysis was not performed. Changes

from baseline in FEV<sub>1</sub> in experimental group and control group were  $0.07 \pm 0.3$  and  $-0.04 \pm 0.2$ , respectively. However, the *P* values were not available.

# 3.7. Subgroup Analysis (AT Adjunctive to Other Treatments versus Other Treatments Alone)

3.7.1. 6MWD. Four studies [9, 16, 21, 25] provided numerical data for 6MWD and were included in the meta-analysis. Analysis of the data indicated that there was heterogeneity ( $\chi^2 = 32.35$ , P < 0.00001;  $I^2 = 91\%$ ); hence, a random-effect model was used. The pooled results showed that 6MWD in the experimental group improved more compared to the control group (MD: 35.15; 95% CI: 2.37 to 67.92; Z = 2.10, P = 0.04) (Figure 13).

3.7.2. SGRQ. Only one study [8] provided numerical data for SGRQ total scores; thus, the meta-analysis was not performed. Compared to the control group (7.0 ± 14.9), both AT plus PR group and PR group demonstrated a significant change for SGRQ total scores ( $-7.1\pm12.7$ , P = 0.01;  $-7.4\pm8.7$ , P = 0.0006). However, there were no data available for symptom domain scores, activity domain scores, and impact domain scores of SGRQ.

Study on Submour	Exp	perimei	ntal	Control			Mainhe	Mean Difference		Mean	Differe	ence	
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI	IV, Random, 95% CI				
Feng et al. 2016	67.4	21.7	33	-13.8	54.8	32	29.7%	81.20 [60.82, 101.58]					
Ge et al. 2017	58.05	51.52	22	19.03	48.18	19	23.3%	39.02 [8.48, 69.56]					
Suzuki et al. 2012	63.5	49.9	30	-19.4	48.7	32	27.0%	82.90 [58.33,107.47]					
Tong et al. 2014	49.23	53.9	16	11.87	48.35	14	20.0%	37.36 [0.77, 73.95]			-	<u> </u>	
Total (95% CI)			101			97	100.0%	63.05 [39.27, 86.83]				◆	
Heterogeneity: $\tau^2 = 387.99$ ; $\chi^2 = 9.17$ , df = 3 ( $P = 0.03$ ); $I^2 = 67\%$										-100	0	100	200
Test for overall effect: $Z = 5.20$ ( $P < 0.00001$ )								F	avours [control]	Favo	ours [experim	ental]	

FIGURE 11: AT adjunctive to other treatments versus placebo or sham acupuncture adjunctive to other treatments, 6MWD.

Study or Subgroup	Exp	erime	ntal	Control			Weight	Mean Difference		Mean	n Diffe	erence	
study of Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI		IV, Ran	dom,	95% CI	
Feng et al. 2016	-15.3	24.8	33	0.5	20	32	28.4%	-15.80 [-26.74, -4.86]			-		
Suzuki et al. 2012	-16	9.7	30	0.3	7.9	32	36.0%	-16.30 [-20.72, -11.88]					
Tong et al. 2014	-4.13	4.77	16	-3.29	8.28	14	35.6%	-0.84 $[-5.77, 4.09]$					
Total (95% CI)			79			78	100.0%	-10.66 [-22.24, 0.92]					
Heterogeneity: $\tau^2 = 91.76$ ; $\chi^2 = 22.16$ , df = 2 ( <i>P</i> < 0.0001); $I^2 = 91\%$										-25	0	25	50
Test for overall effect: $Z = 1.80$ ( $P = 0.07$ )									Favou	rs [experimenta	al]	Favours [control]	

FIGURE 12: AT adjunctive to other treatments versus placebo or sham acupuncture adjunctive to other treatments, SGRQ total scores.

Study or Subgroup	Exp	perimer	ıtal	Control			Waight	Mean Difference		Mean	Differen	се	
study of Subgroup	Mean	SD	Total	Mean	SD	Total	weight	IV, Random, 95% CI		IV, Rano	lom, 95%	6 CI	
Deng et al. 2016	49.4	42.26	22	20.1	46.17	22	23.6%	29.30 [3.15, 55.45]					
Liu et al. 2015	53.1	34.27	40	44.7	30.81	40	26.4%	8.40 [-5.88, 22.68]					
Tang 2017	119.13	42.26	30	34.07	46.17	29	24.6%	85.06 [62.45, 107.67]					
Yang et al. 2009	43.1	42.26	40	23	46.17	40	25.4%	20.10 [0.70, 39.50]					
Total (95% CI)			132			131	100.0%	35.15 [2.37, 67.92]					
Heterogeneity: $\chi^2 = 1004.91$ ; $\chi^2 = 32.35$ , df = 3 ( $P < 0.00001$ ); $I^2 = 91\%$										-100	0	100	200
Test for overall effect: $Z = 2.10$ ( $P = 0.04$ )									Fa	vours [control]	Favou	rs [experime	ental]

FIGURE 13: AT adjunctive to other treatments versus other treatments alone, 6MWD.

3.7.3. *FEV*<sub>1</sub>. Two studies [7, 19] provided numerical data for FEV<sub>1</sub> and were included in the meta-analysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 0.00$ , P = 1.00;  $I^2 = 0\%$ ); hence, a fixed-effect model was used. The pooled results showed that FEV<sub>1</sub> in the experimental group improved more compared to the control group (MD: 0.41; 95% CI: 0.28 to 0.54; Z = 6.01, P < 0.00001) (Figure 14).

3.7.4. Effective Rate. Seven studies [7, 13–16, 19, 25] provided categorical data for effective rate and were included in the meta-analysis. Analysis of the data indicated that heterogeneity was not statistically significant ( $\chi^2 = 2.38$ , P = 0.88;  $I^2 = 0\%$ ); hence, a fixed-effect model was used. The pooled results showed that effective rate in the experimental group was higher compared to the control group (OR: 2.84; 95% CI: 1.59 to 5.06; Z = 3.53, P = 0.0004) (Figure 15).

*3.8. Reporting Biases.* We did not investigate publication biases by funnel plot because each comparison included not more than 10 studies.

### 4. Discussion

This systematic review provided a detailed summary of the current evidences related to the efficacy and safety of AT for functional effects and quality of life in COPD patients.

6MWD is an important measure of functional exercise capacity of patients with COPD. The distance walked is associated with clinical outcomes such as hospitalization and mortality, and its changes are used to evaluate the efficacy of therapeutic interventions such as pulmonary rehabilitation, surgery, and pharmaceutical management [33, 34]. In this review, 6MWD in the experimental group improved more compared to the control group, and the MD was 47.84

Experimental Study or Subgroup Mean SD Total			ntal Total	Mean	Contro SD	l Total	Weight	Mean Difference IV, Fixed, 95% CI		Mean Difference IV, Fixed, 95% CI				
Chu and Cai 2015	0.53	0.43	32	0.12	0.56	32	29.9%	0.41 [0.17, 0.65]						
Wan et al. 2009	0.51	0.43	75	0.1	0.56	75	70.1%	0.41 [0.25, 0.57]						
Total (95% CI)			107			107	100.0%	0.41 [0.28, 0.54]			•			
Heterogeneity: $\chi^2 = 0.00$ , df = 1 (P = 1.00); $I^2 = 0\%$ Test for overall effect: Z = 6.01 (P < 0.00001)										–0.5 Favours [control]	0 0.5 Favours [expe	+ 1 erimental]		

FIGURE 14: AT adjunctive to other treatments versus other treatments alone,  $FEV_1$ .

Study or Subgroup	Experir	nental	Control		Waight	Odds Ratio		O	dds Ratio		
Study of Subgroup	Events	Total	Events	Total	weight	M-H, Fixed, 95% CI		M-H, 1	Fixed, 95% (	CI	
Chu and Cai 2015	30	32	24	32	10.2%	5.00 [0.97, 25.77]				-	
Jia 2004	21	22	17	22	5.3%	6.18 [0.66, 58.03]				-	
Li 2015	26	29	25	30	17.3%	1.73 [0.37, 8.03]		—			
Li et al. 2016	28	30	27	30	12.2%	1.56 [0.24, 10.05]					
Liu et al. 2015	38	40	32	40	10.9%	4.75 [0.94, 23.98]					
Tang 2017	26	30	22	29	20.3%	2.07 [0.53, 8.00]					
Wan et al. 2009	71	75	66	75	23.9%	2.42 [0.71, 8.24]					
Total (95% CI)		258		258	100.0%	2.84 [1.59, 5.06]				•	
Total events	240		213								
Heterogeneity: $\chi^2 = 2$	$(38); I^2 = 0$	0.01	0.1	1	10	100					
Test for overall effect: $Z = 3.53$ ( $P = 0.0004$ )								Favours [control]	Favour	s [experim	ental]

FIGURE 15: AT adjunctive to other treatments versus other treatments alone, effective rate.

meters, which was greater than 25 meters, the minimal clinically important difference (MCID) of 6MWD for COPD patients [34]. This result might indicate the potential of AT in improving exercise capacity of COPD patients. Two subgroup analyses supported this result as well, and the MD of 6MWD change was 63.05 meters and 35.15 meters, respectively.

SGRQ, another primary outcome measure in this review, is a well-established disease-specific instrument to measure quality of life for asthma and COPD. In this review, there was no statistically significant improvement in SGRQ total scores between two groups. However, MD of symptom domain scores, activity domain scores, and impact domain scores of SGRQ was 24.86 units, 16.52 units, and 13.07 units, respectively. Although there was no MCID available for each domain, each MD was at least three times greater than 4 units, the MCID for SGRQ total scores in COPD patients [35], and this might suggest the effect of AT on different aspects of health status in COPD patients. Subgroup analysis (AT adjunctive to other treatments versus placebo or sham acupuncture adjunctive to other treatments) supported these above results as well.

 $FEV_1$  is widely used by physicians in the diagnosis, classification, treatment, monitoring, and establishing prognosis for COPD patients. In this review, there was no statistically significant improvement in  $FEV_1$  between two groups. However, subgroup analysis (AT adjunctive to other treatments versus other treatments alone) showed MD of  $FEV_1$  change was 410 mL, which was four times greater than 100 mL, the MCID of  $FEV_1$  for COPD patients [36]. And this result might

suggest the potential of AT in improving pulmonary function in COPD patients.

mMRC is a major instrument to measure breathlessness. In this review, since mMRC scores were only available in one study, the meta-analysis was not performed. According to this study, change from baseline in mMRC scores in AT plus PR group and PR group was 0.3 units in both, and it was reported that there was significant difference within AT plus PR group. However, it was limited to support the effect of AT in improving breathlessness in COPD patients.

Effective rate, an important outcome measure in clinical studies of Chinese medicine, was also evaluated. In this review, effective rate in the experimental group was higher compared to the control group; to some extent, this might suggest that AT was a more effective treatment compared to other treatments. Importantly, subgroup analysis (AT adjunctive to other treatments versus other treatments alone) also supported this result with OR of 2.84.

Adverse effects were poorly reported in included studies. One study reported some minor adverse effects, and 5 studies reported no adverse effects. This might indicate the safety of AT for COPD patients.

There were some limitations in this study. Firstly, methodological quality of the included studies was generally low. For example, most of the included studies had high risk of performance bias. Secondly, most analysis of the data in the meta-analysis indicated that there was heterogeneity. Thirdly, there were various intervention forms of AT, which might make it difficult to evaluate the efficacy of AT alone. Finally, some resources with language other than English and Chinese might not be included in this review.

### 5. Conclusions

AT may be effective and safe in improving functional effects and quality of life in COPD patients. Besides, AT may also improve pulmonary function of COPD patients. Evidences are inadequate to support the potential of AT in improving breathlessness of COPD patients. These evidences may be useful to clinicians, patients, and health policy-makers with regard to application of AT in COPD. However, further highquality RCTs are needed to confirm the efficacy and safety of AT for COPD patients.

### **Additional Points**

*Registration Number.* This article is registered with PROSPERO 2016 CRD42016054335 (available from http://www.crd.york.ac.uk/PROSPERO/display\_record.php?ID= CRD42016054335).

## **Conflicts of Interest**

The authors declare that there are no conflicts of interest.

## **Authors' Contributions**

Jiajia Wang searched the literature, conducted the statistical analysis, and drafted the manuscript. Yang Xie and Xueqing Yu screened the studies, extracted the data, and evaluated the risk of bias. Jiansheng Li revised the manuscript. Yang Xie conceived this study.

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# **Supplementary Materials**

The methods for this study had been developed according to the recommendations of the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) statement. (Supplementary Materials)

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