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Research Article

Acupuncture and Related Therapies for Obesity: A Network Meta-Analysis

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Obesity is a worldwide public health problem. Currently, increasing evidence suggests acupuncture and related therapies are effective for obesity. This network meta-analysis (NMA) was performed to compare the effectiveness of different acupuncture and related therapies. We searched potential randomized controlled trials (RCTs) in three international databases. Thirty-four trials involving 2283 participants were included. Pairwise meta-analysis showed that acupuncture and related therapies were superior to lifestyle modification and placebo in reducing weight and body mass index (BMI). Based on decreases in body weight, results from NMA showed that acupoint catgut embedding (standard mean difference [SMD]: 1.26; 95% credible interval [95% CI], 0.46–2.06), acupuncture (SMD: 2.72; 95% CrI, 0.06–5.29), and combination of acupuncture and related theories (SMD: 3.65; 95% CI, 0.96–6.94) were more effective than placebo. Another NMA result indicated that acupoint catgut embedding (SMD: 0.63; 95% CI, 0.25–1.11), acupuncture (SMD: 1.28; 95% CrI, 0.43–2.06), combination of acupuncture and related therapies (SMD: 1.44; 95% CrI, 0.64–2.38), and electroacupuncture (SMD: 0.60; 95% CrI, 0.03–1.22) were superior to lifestyle modification in decreasing BMI. Combination of acupuncture and related therapies was ranked the optimal method for both reducing weight and BMI. Further studies will clarify which combination of acupuncture and related therapies is better.

1. Introduction

Obesity, a worldwide public health problem, is described as an adiposity-based chronic disease [1]. Currently, guidelines recommended using body mass index (BMI) to classify individuals as having obesity (BMI ≥30 kg/m²) [2]. Based on the survey conducted previously, the standardized prevalence rates for obesity in adult were 34.9% in United States and 17.7% in China [3, 4]. Moreover, it is associated with other health concerns, such as insulin resistance, type 2 diabetes

mellitus, cardiovascular disease, and cancer, which increased individuals and societies' medical burden [5].

Lifestyle modification, pharmacotherapy, and bariatric surgery are considered the mainstay of therapy for obesity [2]. Although diet and exercise play an essential role in the weight management, their precise mode of action remains controversial [6]. Five long-term medicines (naltrexone-bupropion, phentermine-topiramate orlistat, lorcaserin, and liraglutide) have been approved by US Food and Drug Administration (FDA) for the treatment of obesity [7]. The

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latest research suggested that phentermine-topiramate was associated with the highest possibility of achieving at least 5% weight loss [7]. However, little is known about the long-term safety profile of pharmacotherapy for weight loss. The effectiveness of bariatric procedures for treating obesity has been reported in several randomized controlled trials (RCTs) [8–10]. Nevertheless, the evidence on cardiovascular disease and mortality remains to be validated [11]. Therefore, it is necessary to explore other forms of alternative therapies which are both safe and effective in preventing gaining weight.

In reviewing the literature, it became evident that acupuncture and related therapies have been wildly used for obesity treatment. As mentioned in the meta-analysis, combination of acupuncture and lifestyle modification is more effective compared with lifestyle modification alone [12]. Results of Yeh's research suggested that ear acupoint stimulation had remarkable improvements in the anthropometric parameters of Body Weight (BW), BMI, and so on [13]. In addition, another systematic review performed in 2015 has also shown that clinical efficacy of acupoint catgut embedding therapy was better than that of the control group for simple obesity [14]. However, a major problem is that whether acupuncture or acupuncture-related therapies alone or combined therapy is more effective than lifestyle modification management remains disputable.

By using the technique of network meta-analysis (NMA), both direct and indirect randomized data can be analyzed, and recommended rankings of different treatments can be provided [15, 16]. Therefore, we conducted this Bayesian network meta-analysis to analyse both direct and indirect comparisons of acupuncture and related methods for treating obesity. In this paper, changes in BW, BMI, and the rates of complications of included studies were analyzed.

2. Methods

Our research was conducted following the Preferred Reporting Items for Systematic Reviews and Meta-Analyses for Network Meta-Analysis (PRISMA-NMA) checklist [17] (see Appendix 1).

2.1. Data Sources and Search Strategy. Three electronic international databases (PubMed/Medline, Embase, and the Cochrane Library) were searched for potential RCTs (randomized controlled trials). We identified articles published from initiation to December 2017 with a limit to studies of RCT and without limitations on language or the form they are

published in. The complete search strategies are shown in Appendix 2.

2.2. Study Selection. Two researchers (XC and HL) independently identified irrelevant research based on titles and abstracts. Additionally, full-text articles were scanned by these two researchers to identify eligible studies. All disagreements were resolved by consensus and adjudged by a third reviewer (TC) if necessary. In case of duplicate citations, the most updated studies were selected for data extraction.

2.3. Inclusion and Exclusion Criteria. The studies included in the NMA met the following criteria: (1) the study design must be a randomized controlled clinical trial (RCT); (2) patients diagnosed with simple obesity irrespective of ages and sex as study subjects; diagnostic criteria must be clear and inclusion and exclusion criteria were explicit; (3) at least one of the following efficacy outcomes or safety endpoints was included: BW, BMI, and adverse events; (4) participants in the experimental group have received acupuncture and related treatments (specifically, classical body acupuncture; electroacupuncture auricular acupoint stimulation; acupoint catgut embedding and warming acupuncture) alone or in combination; (5) English or Chinese language studies.

The following were excluded: (1) self-control and non-RCTs; (2) preclinical studies, systematic reviews, case reports, and meta-analyses; (3) reports without sufficient and clear original data; (4) participants having received other forms of acupuncture such as transcutaneous electrical nerve stimulation or laser acupuncture; (5) duplicate studies and studies reporting the same results.

2.4. Data Collection and Quality Assessment. According to a standard data collection sheet, two investigators (TC and XZ) independently extracted the following data: (1) main characteristics of included randomized controlled trials (i.e., year of publication, type of intervention, patients characteristics, types of outcome, and reported adverse events); (2) details of acupuncture and related interventions (i.e., frequency and duration of acupuncture sessions, names of acupuncture points used, and retention time); (3) clinical outcome (i.e., summaries of mean, standard difference, and sample size between treatment groups). In some trials, the change between baseline and after treatment was failed to present. Using the methods recommended in the Cochrane Handbook for Systematic Reviews of Interventions (version 5.1) [18], the missing data was estimated using the following formula:

$$\overline{X}_{change} = \overline{X}_{post-treatment} - \overline{X}_{baseline}$$
 (1)

$$SD_{change} = \sqrt{\left(SD_{baseline}\right)^2 + \left(SD_{post-treatment}\right)^2 - 2 \times r \times SD_{baseline} \times SD_{post-treatment}}$$
 (2)

where r is a correlation coefficient with a value of 0.5 [19]. For each included RCT, two researches (XT and XC)

independently assessed their risk of bias by the Cochrane Collaboration tool [20]. Bias risks of each study were assessed

from six aspects: random sequence generation, allocation concealment, blinding of participants and investigators, blinding of outcome assessment, incomplete outcome data addressed, and selective outcome reporting, while ranked in high risk, low risk, and unclear risk.

2.5. Statistical Analysis. Firstly, standard pairwise meta-analysis was initially performed using the Review Manager (Version 5.3, Cochrane Collaboration, Oxford, UK). We calculated I-square ($\rm I^2$) test to assess heterogeneity among RCTs [21]. To be specific, when there was $\rm I^2 > 50\%$, they were analysed using a random effects model; otherwise, a fixed effect model was chosen. Subgroup analyses were conducted according to the type of acupuncture treatment and the treatment of control group. Mean difference (MD) with 95% confidence intervals (CI) was used to analyze continuous data. We generated forest plots to illustrate the relative strength of curative effects.

Second, to indirectly compare the effectiveness among treatments of acupuncture and related therapies, we did a random effects model NMA within a Bayesian framework, by using WinBUGS (Version 1.4.3, MRC Biostatistics Unit, Cambridge, UK) [22, 23]. Models were computed with Markov chain Monte Carlo (MCMC) simulation methods, using four chains with overdispersed initial values. We utilized the Markov chains for 50,000 simultaneous iterations after the first 20000 iterations were discarded because they may have an influence on the arbitrary value. In this process, the convergence of the model was assessed by the Brooks-Gelman-Rubin (BGR) method; a value of potential scale reduction factor (PSRF) close to 1 indicated the better convergence [24]. The continuous outcome was measured by a standard mean difference (SMD) with a 95% credible intervals (CrI) for indirect comparisons.

Finally, plot of surface under the cumulative ranking curve (SUCRA) was generated using the STATA software (Version 13.0; Stata Corporation, College Station, Texas, USA), which indicated the probability of each intervention of being ranked best [25]. In our study, higher SUCRA scores mean the higher rank of the treatment [15]. A Z value and its corresponding p-value were calculated, and an R value less than 0.05 indicated a statistically significant difference.

3. Results

- 3.1. Study Search. We performed this research on Dec 26 2017. As shown in Figure 1, a total of 1050 records were initially identified from the databases. 675 studies left after duplicates were removed. 577 records were excluded after carefully scanning titles and abstracts. Finally, 34 trials with 2283 participants were included in our NMA [26–59], covering 8 groups, manual acupuncture; electroacupuncture; auricular acupoint stimulation; acupoint catgut embedding; pharmacotherapy; warming acupuncture; lifestyle modification; placebo.
- 3.2. Study Description. Main characteristics of included RCTs were shown in Table 1. The participants were from Australia [28], the United States [26], Turkey [46], Korea [51], Iran

[36], Egypt [48], and China. Age of participants ranged from 15 to 70 years, while the sample size of the studies ranged from 12 to 86. Among the included RCTs, there were one four-arm trials, 5 three-arm trials, and 28 two-arm trials. Fourteen studies compared acupuncture to placebo. Ten studies compared acupuncture to lifestyle intervention. Six studies compared combined therapies to acupuncture alone. Details about acupuncture points used, retention time, frequency, and duration of acupuncture sessions were shown in Table 2. In these research, 30 articles [26-30, 32, 34-38, 41-59] reported the weight loss, while 25 articles reported the change in BMI. The details of mean, standard difference (SD), and sample size between different groups for eligible studies are summarized in Appendix 3. The Cochrane risk of bias assessment was presented in Table 3. Furthermore, the network plot of included comparisons was shown in Figure 2.

3.3. Pairwise Meta-Analyses

3.3.1. Body Weight. A direct pairwise meta-analysis showed that acupuncture and related therapies showed a greater BW reduction than lifestyle modification (MD: 1.66; 95% Confidence interval, 0.63to2.70) and placebo (MD: 1.15; 95% CI, 0.67to1.63). When compared to acupuncture, combination of acupuncture and related theories showed a marginally stronger effect in weight loss (MD: 1.56; 95% CI, 0.07to3.05). There was no statistically significant difference between combination of acupuncture and related theories and pharmacotherapy in their effectiveness in BW (MD: 2.44; 95% CI, -1.98to6.86). (Table 4)

BMI. As for the comparison in reducing BMI, acupuncture and related therapies were found to be marginally superior to lifestyle modification (MD: 1.17; 95% CI, 0.09to2.26) and placebo (MD: 0.57; 95% CI, 0.40to0.74). The remaining direct comparisons did not show significant differences (Table 4).

3.4. Network Meta-Analysis

3.4.1. Body Weight. The NMA showed that all treatments other than acupuncture combined lifestyle modification were more efficacious than lifestyle modification. Three treatments were significantly more effective than placebo. Specifically, acupoint catgut embedding (SMD: 1.26; 95% credible interval, 0.46to2.06), acupuncture (SMD: 2.72; 95% CrI, 0.06to5.29), and combination of acupuncture and related therapies (SMD: 3.65; 95% CrI, 0.96to6.94). Furthermore, moxibustion with warming needle was associated with a significantly improvement than lifestyle modification (SMD: -5.24; 95% CrI, -10.15to-0.55) (Table 5).

3.4.2. BMI. Four treatments showed superiority over placebo, including acupoint catgut embedding (SMD: 1.31; 95% CrI, 0.36to2.06), acupuncture (SMD: 1.94; 95% CrI, 0.83to3.00), combination of acupuncture and related theories (SMD: 3.65; 95% CrI, 0.96to6.94), and electroacupuncture (SMD: 1.28; 95% CrI, 0.43to2.11). Four treatments were significantly more effective than lifestyle modification, including acupoint catgut embedding (SMD: 0.63; 95% CI,

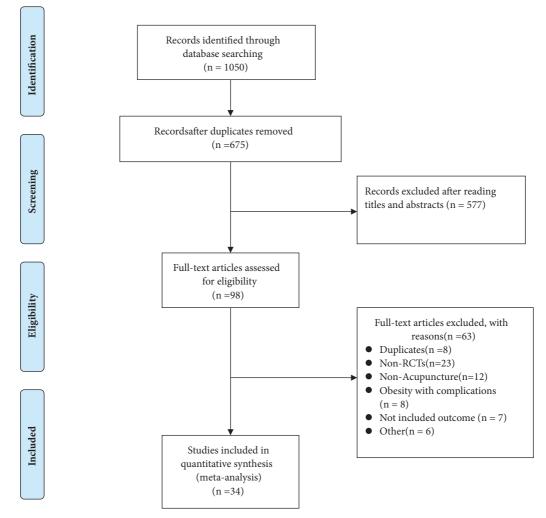


FIGURE 1: PRISMA flow chart.

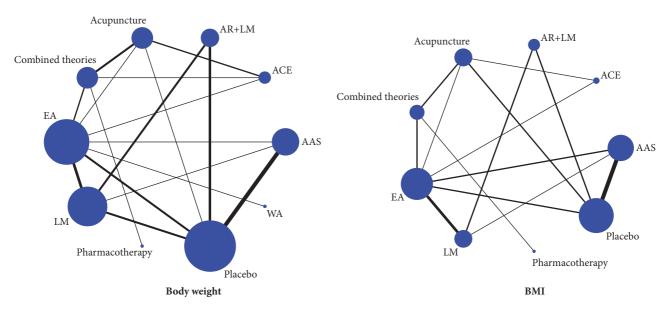


FIGURE 2: Network plot. BMI: body mass index; LM: lifestyle modification; AAS: auricular acupoint stimulation; EA: electroacupuncture; ACE: acupoint catgut embedding; WA: warming acupuncture; AR: acupuncture and related therapies; combined therapies: combination of acupuncture and related therapies.

Table 1: Main characteristics of included randomized controlled trials.

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Study ID and Country	Sample size R/A	Age: mean (SD) or range R/A	Intervention	Control	Adverse events reported R/A	Type of outcomes
Allison et al. [17] 1995, USA	35/34	19 - 70	AAS	Placebo	Redness, pain, bleeding	BW,
Hsu et al. [18] 2005, Taiwan	24/22	41.5(11.2)/41.0 (10.0)	EA	LM	Ecchymosis(2), abdominal discomfort(1)/None	BW, BMI
Richards et al. [19] 1998, Australia	28/32	44.1 (11.7)/43.0 (13.6)	AAS	Placebo	intercurrent illness and discontinued(1)/None	BW
He et al. [20] 2008, China	40/40	18 - 50	Combined therapies#	Pharmacotherapy	NR	BW, BMI
Li et al. [21] 2006, China	26/30 29/30	16.00(1.38)/16.00(1.95) 15.00(2. 04)/16.00(1.95)	EA AAS	LM	NR	BMI
Tong et al. [22] 2011, China	76/42	35.08(9.31)/34.60(8.55)	Acupuncture	Placebo	Adverse events VAS	BMI
Hsu et al. [23] 2009, Taiwan	23/22	40.0 (10.5)/39.4 (13.6)	AAS	Placebo	Minor-inflammation(1), mild tenderness (7)/mild tenderness (2)	BW, BMI
Hsieh et al. [24] 2010, Taiwan	26/26	18 - 20	AAS	Placebo	NR	BMI
Hsieh et al. [25] 2011, Taiwan	27/28	18 - 20	AAS	Placebo	NR	BW
Abdi et al. [26] 2012, Iran	86/83	37.29(1.0)/38.73 (1.1)	AAS	Placebo	None	BW, BMI
Darbandi et al. [27] 2012, Iran	43/43	37.57(9.26)/37.65(9.71)	AR+ LM	Placebo	None	BW, BMI,
He et al. [28] 2012, China	30/30	18-54	AR+ LM	TM	NR	BW, BMI
Lien et al. [29] 2012, Taiwan	24/23	39.2(11.6)/40.7 (9.7)	AAS	Placebo	Dizziness (1)/None	BW, BMI
Darbandi et al. [30] 2014, Iran	20/20 20/20	38.0(0.9)/38.0(1.3) 39.0(1.8)/37.9(1.5)	EA AAS	Placebo(EA) Placebo(AAS)	None	BW, BMI
Yeh et al. [31] 2015, Taiwan	36/34	29.9 (7.7)/32.8 (9.5)	EA	Placebo	NR	BMI
Chen et al. [32] 2007, China	40/40	43.1(13.6)/44.6(10.3)	ACE	Acupuncture	NR	BW, BMI
Huang et al. [33] 2011, China	30/30	NR	ACE	EA	NR	BW, BMI
Tang et al. [34] 2009, China	33/32	21-54/22-55	Combined therapies	EA	NR	BW, BMI

TABLE 1: Continued.

		TYPET	rable i. Commuca.			
Study ID and Country	Sample size R/A	Age: mean (SD) or range R/A	Intervention	Control	Adverse events reported R/A	Type of outcomes
Shi et al, [35] 2006, China	40/42	17-49/18-51	Combined therapies	EA	NR	BW, BMI
Hsu et al. [36] 2005, Taiwan	22/20	40.0 (11.5)/41.3 (9.9)	EA	LM	mild Ecchymosis(3), abdominal	BW, BMI
					discomfort(1)/None	
Güçel et al. [37] 2012, Turkey	20/20	$34.6\pm6.3/36.8\pm7.8$	Acupuncture	Placebo	NR	BW, BMI
Deng et al. [38] 2014, China	30/30 30/30	32(7)/33(7) 32(7)/33(8)	Combined therapies	Acupuncture ACE	NR	BW
Hassan et al. [39] 2014, Egypt	21/30	45.00 (9.32)/43.47 (9.59)	AR+ LM	ΓM	NR	BW, BMI
He et al. [40] 2014, China	28/28	NR	Combined therapies	Acupuncture	NR	BW, BMI
Wang et al. [41] 2013, China	45/45	31(10)/32(12)	EA	Acupuncture	NR	BMI
Sujung et al. [42] 2014, South Korea	22/15	34.7(11.9)/42.7(10.2)	AAS	Placebo	NR	BW, BMI
Bu et al. [43] 2007, China	32/23	32.1(1.1)/33.4(1.3)	Combined therapies	Acupuncture	NR	BW, BMI
Shi et al. [44] 2005, China	36/32	19~58/18~56	WA	EA	NR	BW
Yang et al. [45] 2010, China	31/30	18~42/18~48	AR+ LM	ΓW	NR	BW
Cabioglu et al. [46] 2005, Turkey	22/12 22/21	39.8(5.3)/43.3(4.3) 39.8(5.3)/42.7(3.9)	EA	Placebo LM	NR	BW
Cabioglu et al. [47] 2006, Turkey	20/15 20/15	42.1(4.4)/41.8(4.6) 42.1(4.4)/42.9 (4.3)	EA	Placebo LM	NR	BW
Cabioglu et al. [48] 2008, Turkey	20/15 20/23	40.55 (5.30)/41.47 (4.61) 40.55 (5.30)/42.91(4.02)	EA	Placebo LM	NR	BW
Darbandi et al. [49] 2013, Iran	42/44	36.50 (9.26)/36.48 (8.69)	AR+ LM	Placebo	NR	BW, BMI
Fogarty et al. [50] 2015, Australia	19/16	>18	AR+ LM	Placebo	NR	BMI

BW: body weight; BMI: body mass index; LM: lifestyle modification; AAS: auricular acupoint stimulation; EA: electroacupuncture; ACE: acupoint catgut embedding; WA: warming acupuncture; AR: acupuncture and related therapies.

TABLE 2: Descriptions of the included acupuncture and related therapies.

Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Allison et al. 1995, USA	AAS	NR	2-3 min	3 sessions daily for 12 weeks
Hsu et al. 2005, Taiwan	EA	Qiai(REN9), Shuifen(REN9) Shuidao(ST28), Siman(K14) Zusanli(ST26), Fenglong(ST40) Sanginjao(SP6)	40 min	2 sessions weekly for 6 weeks
Richards et al. 1998, Australia	AAS	Shenmen(TF4), Stomach(CO4)	15-20 min	2 sessions daily for 4 weeks
He et al. 2008, China	Combined therapies#	Ear acupressure: Shenmen(TF4), Neifenmi(CO18), Pi(CO13), Wei(CO14), Sanjiao(CO17), Dachang(CO7), Naodian Body acupuncture: Tianshu(ST25), Guanyuan(RN4) Sanyinjiso(SP9), Fenglong(ST40) Zusanli(ST36)	Ear acupressure:3 days Body acupuncture:30 min	Ear acupressure:1 session every 3 days with a total of 10 sessions Body acupuncture: The first 5 days of treatment 1 time, 5 days after treatment 1, 1 month, for a course of treatment.
1; of al 2008 (China	EA	Sanginjao(SP6), Tianshu(ST25) Zusanli(ST36), Quchi(LIII) Fenglong(ST40), Neiting(ST44) Zhongwan(CV12), Pishu(BL20) Shenshu(BL23), Qihai(CV6) Yinlingquan(SP9), Shangjuxu(ST37) Taichong(LR3)	10 min	l session daily with a total of 60 sessions, 2 days rest in-between 10 sessions
2000, Cillia	AAS	Hunger point Pizhixia(AT4) Shenmen(TF4), Shenshangxian(TG2P) Sanjiao(CO17), Pi(CO13) Wei(CO14), Fei(CO14) Kou(CO1), Dachang(CO7) Zhichangxiduan(HX2)	15-20 min	1 session daily with a total of 10 sessions for 10 weeks, 2 day rest in-between 10 sessions
Tong et al. 2011, China	Acupuncture	Zhongwan(CV12), Zhongji(CV3) Daheng(SP15), Xiawan(CV10) Shimen(CV5), Tianshu(ST25) Liangqiu(ST34), Zusanli(ST36) Yinlingquan(SP9)	30 min	1 session every other day for a total of 5 weeks with 12 sessions
Hsu et al. 2009, Taiwan	AAS	Hunger point, Shenmen point(TF4) Stomach point(CO4), Endocrine point(CO18)	3 days	2 sessions weekly for a total of 6 weeks with 12 sessions
Hsieh et al. 2010, Taiwan	AAS	NR	2/3 days	1 session weekly for 10 sessions
Hsieh et al. 2011, Taiwan	AAS	NR	NR	l session weekly for a total of eight weeks

TABLE 2: Continued.

Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Abdi et al. 2012, Iran	AAS	Shenmen(TF4), Stomach(CO4) Hunger point Mouth(CO1) Centre of ear(HXI), Sanjiao(CO17)	3 days	Twice a week for a total of 6 weeks
Darbandi et al. 2012, Iran	AAS	Shenmen(TF4), Stomach(CO4) Hunger point Mouth(CO1) Centreof ear(HX1), Sanjiao(CO17)	3 days	Twice a week for a total of 6 weeks
He et al. 2012, China	AAS	Hunger point Stomach(CO4) Spleen(CO14), LargeIntestine(CO7) Endocrine(CO18), Shenmen(TF4)	3 days	3 times a day for 4 weeks
Lien et al. 2012, Taiwan	AAS Placebo	Shenmen point(TF4), Stomach point(CO4) Hunger point, Endocrine point(CO18) Shenmen point(TF4), Stomach point(CO4)	N N NR	3 session weekly with a total of 12 sessions for 4 weeks 3 sessions weekly with a total of 12
		point(CO18)		
Darbandi et al. 2014, Iran	EA	Tianshu (ST-25), Weidao(GB28) Zhongwan(REN12), Shuifen(REN9) Guanyuan(REN4), Sanyinjiao(SP6) Quchi(LIII), Fenlong(ST40) Qihai(REN6), Yinlingquan (SP9) Shenmen (TF4), Stomach (CO4) Hunger point, Mouth (CO1) Center of ear (HX1), Sanjiao (CO17)	20 min 3 days	2 sessions weekly for a total of 6 weeks 2 sessions weekly for a total of 6 weeks
Yeh et al. 2015, Taiwan	EA	Shenmen (TF4), Stomach CO4) Endocrine (CO18) Hunger point	20 min	NR
7007 Point	ACE	Liangqiu(ST34), Zhongwan(CV12) Tianshu(ST25), Shuifen(CV9) Fenglong(ST40)	A week	1 session weekly with a total of 30 sessions for 4 weeks
Cnen et al. 2007, Cnina	Acupuncture	Liangqiu(ST34), Zhongwan(CV12) Tianshu(ST25), Shuifen(CV9) Fenglong(ST40)	45 min	The first 5 days are 1 times a day, and 1 time after 5 days, 1 month is 1 course of treatment.
	ACE	One set is Tianshu(ST25) Zhongwan(CV12), Guanyuan(CV4) Zusanli(ST36), Weishu(BL21) Ashixue	24 hour	1 session weekly with a total of 7 sessions for 60 days
Huang et al. 2011, China	EA	Zhongwan(CV12), Tianshu(ST25) Daheng(SP15), Shuifen(CV9) Qlhai(CV6), Guanyuan(CV4) Zusanli(ST36), Ashixue	30 min	3 sessions weekly with a total of 12 sessions for 60 days

TABLE 2: Continued.

		TABLE 2: Continued.		
Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Tang et al. 2009, China	Combined therapies	EA: Zhongwan(CV12), Xiawan(CV10) Guanyuan(CV4), Tianshu(ST25) Shuifen(CV9), Sanyinjiao(SP6) Zusanli(ST36), Xuehai(SP10) Xinshu(BL15), Geshu(BL17) Pishu(BL20) ACE: Zhongwan(CV12), Tianshu(ST25) Qihai(CV6), Tianshu(ST25) Liangqiu(ST34), Zusanli(ST36) Gongsun(SP4), Xinshu(BL15) Pishu(BL20) Zhongwan(CV12), Xiawan(CV10)	EA:30 min	EA:The first 3 days are 1 times a day, and 1 time after 3 days, 15 days is 1 course of treatment. ACE:After the first acupoint catgut embedding for 3 consecutive times, the interval is buried for the second time after 15 days, and the acupuncture is performed for the third time after the end of the treatment period.
	EA	Guanyuan(CV4), Tianshu(S125) Shuifen(CV9), Sanyinjiao(SP6) Zusanli(ST36), Xuehai(SP10) Xinshu(BL15), Geshu(BL17) Pishu(BL20)	30 min	The first 3 days are 1 times a day, and 1 time after 3 days, 15 days is 1 course of treatment.
Shi et al. 2006, China	Combined therapies	Zhongwan(CV12), Xiawan(CV10) Qihai(CV6), Zhongji(CV3) Tianshu(ST25), Daheng(SPI5) Liangmen(ST21), Huaroumen(ST24) Shuidao(ST28), Quchi(CV6) Zhigou(TE6), Hegu(L14) Liangqiu(ST34), Zusanlli(ST36) Shangjuxu(ST37), Fenglong(ST40) Sanyinjiao(SP6), Gongsun(SP4) Neiting((ST44) Zhongwan(CV12), Xiawan(CV10) Qihai(CV6), Zhongji(CV3) Tianshu(ST25), Daheng(SPI5)	30 min	EA:The first 3 days are 1 times a day, and 1 time after 3 days, 15 days is 1 course of treatment. ACE: After the first acupoint catgut embedding for 3 consecutive times, the interval is buried for the second time after 15 days, and the acupuncture is performed for the third time after the end of the treatment period.
	EA	Liangmen(S121), Huaroumen(ST24) Shuidao(ST28), Quchi(CV6) Zhigou(TE6), Hegu(L14) Liangqiu(ST34), Zusanlli(ST36) Shangjuxu(ST37), Fenglong(ST40) Sanyinjiao(SP6), Gongsun(SP4) Neiting((ST44)	30 min	The first 3 days are 1 times a day, and 1 time after 3 days, 15 days is 1 course of treatment.
Hsu et al. 2005, Taiwan	EA	Qihai (REN-6), Shuifen (REN-9) Shuidao (ST-28), Siman (K-14) Zusanli (ST-26), Fenglong(ST-40) Sanginjao (SP-6)	40 min	2 sessions weekly with a total of 12 sessions for 6 weeks

ABLE 2: Continued.

Subject of acupuncture Names of acupuncture points used Recursion time Frequency & Acupuncture sessions Güçel et al. 2012, Turkey Acupuncture Zheagul (14), Shamazur (1474) 20 min 2 sessions weekly with a total of 2 sessions for 5 weeks assistant of 2 sessions for 5 weeks a session for 5 weeks a sessio			TABLE 2: Commucu.		
Hegul.(14), Sheumen(HT7) Acupuncture	Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Combined therapies	Güçel et al. 2012, Turkey	Acupuncture	Hegu(LI4), Shenmen(HT7) Zusanli(ST36), Neiting(ST44) Sanyinjiao(SP6)	20 min	2 sessions weekly with a total of 10 sessions for 5 weeks
Chongwan (CV 12), Xiawan (CV 10) Chongwan (CV 12), Xiawan (CV 10) Chongwan (CV 12), Wailing (ST 10) Chongwan (CV 12), Tianshu (ST 25) Chanyuan (CV 12), Zhagou (TE 6) Chanyuan (ST 25), Liangmen (ST 21) Chanyuan (ST 25), Liangmen (ST 21) Changwan (ST 25), Liangmen (ST 21) Changwan (ST 25), Liangmen (ST 21) Changwan (ST 25), Liangmen (ST 25) Changwan (ST 25)		Combined therapies	Zhongwan (CV 12), Xiawan(CV 10) Qihai(CV 6), Guanyuan(CV4) Huaroumen (ST 24), Wailing (ST 26) Daheng (SP 15), Tianshu (ST 25) Yinjiao (CV 7), Zhigou (TE 6) Zusanll (ST 36)	NR	Acupuncture:1 session every 3 days with a total of 21 sessions for 4 weeks, 3 days rest between every session Acupoint catgut Embedding:1 session weekly with a total of 3 sessions for 3 weeks
Acupoint catgut embedding	Deng et al. 2014, China	Acupuncture	Zhongwan (CV 12), Xiawan(CV 10) Qihai(CV 6), Guanyuan(CV4) Huaroumen (ST 24), Wailing (ST 26) Daheng (SP 15), Tianshu (ST 25)	30 min	1 session every 3 days with a total of 21 sessions for 4 weeks, 3 days rest between every session
AR AR Tianshu (ST25), Liangmen(ST21) Daheng (SP15), Zusanli (ST36) acupuncture Sanyinjiao(SP6), Quchi (LIII) Zhigou (SJ6), Zhongwan(RN12) Qihai (RN06) Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Tianshu(ST25), Zusanli(ST36) Quchi(LIII) Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Acupuncture Tianshu(ST25), Zusanli(ST36) Quchi(LIII) Shen-men(TF4), Stomach(CO4) AAS Spleen(CO13), Hunger point Endocrine(C018)		Acupoint catgut embedding	Zhongwan (CV 12), Tianshu (ST 25) Yinjiao (CV 7), Zhigou (TE 6) Guanyuan (CV 4), Zusanli (ST 36)	NR	1 session weekly with a total of 3 sessions for 3 weeks
AR Tianshu (ST25), Liangmen(ST21) Daheng (SP15), Zusanli (ST36) acupuncture Sanyinjiao(SP6), Quchi (LIII) Zhigou (SJ6), Zhongwan(RN12) Qihai (RN06) Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Shen-men(TF4), Stomach(CO4) AAAS Spleen(CO13), Hunger point Endocrine(C018)	Hassan et al. 2014, Egypt	AR	NR	NR	NR
Tianshu (ST25), Liangmen(ST21) Daheng (SP15), Zusanli (ST36) Sanyinjiao(SP6), Quchi (LIII) 30 min Zhigou (SJ6), Zhongwan(RN12) Qihai (RN06) Qihai (RN06) Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) 30 min Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Tianshu(ST25), Zusanli(ST36) Acupuncture Shen-men(TF4), Stomach(CO4) Shen-men(TF4), Stomach(CO4) Shen-men(TF4), Hunger point NR Endocrine(C018) Endocrine(AR	NR	NR	NR
et al. 2013, China Acupuncture Acupuncture \$\text{Actan} \text{Neiting(ST44)} \text{Stannli(ST36)} \text{Stomanli(ST36)} \text{30 min} \text{Actannli(ST36)} \text{Actannli(ST36)} \text{30 min} \text{Vainxu(ST37)} \text{Kiajuxu(ST37)} \text{Kiajuxu(ST37)} \text{Kiajuxu(ST37)} \text{Kiajuxu(ST36)} \text{Quchi(LIII)} \text{Quchi(LIII)} \text{Shen-men(TF4), Stomach(CO4)} \text{Shen-men(TF4), Stomach(CO4)} \text{NR} \text{Spleen(CO13), Hunger point} \text{NR} \text{Endocrine(C018)} \text{Randocrine(C018)}	He et al. 2014, China	acupuncture	Tianshu (ST25), Liangmen(ST21) Daheng (SP15), Zusanli (ST36) Sanyinjiao(SP6), Quchi (LII1) Zhigou (SJ6), Zhongwan(RN12) Qihai (RN06)	30 min	1 session daily with a total of 21 sessions for 3 weeks
et al. 2013, China Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Xiajuxu(ST39), Fenglong(ST40) Tianshu(ST25), Zusanli(ST36) Quchi(LIII) Shen-men(TF4), Stomach(CO4) Spleen(CO13), Hunger point Endocrine(C018)	. [0 0000]	EA	Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Tianshu(ST25), Zusanli(ST36) Quchi(LII1)	30 min	1 session every 2 days with a total of 12 sessions for 3 weeks
Shen-men(TF4), Stomach(CO4) Spleen(CO13), Hunger point Endocrine(C018)	Wang et al. 2013, Cnina	Acupuncture	Neiting(ST44), Shangjuxu(ST37) Xiajuxu(ST39), Fenglong(ST40) Tianshu(ST25), Zusanli(ST36) Quchi(LII1)	30 min	1 session every 2 days with a total of 12 sessions for 3 weeks
	Sujung et al. 2014, South Korea	AAS	Shen-men(TF4), Stomach(CO4) Spleen(CO13), Hunger point Endocrine(C018)	NR	1 session weekly with a total of 8 sessions for 8 weeks

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Contin
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TABLE

		TABLE 2: Continued.		
Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Bu et al. 2007, China	Combined therapies	Acupuncture: Tianshu(ST25), Guanyuan(CV4) Zusanli(ST36), Fenglong(ST40) Sanyinjiao(SP6) AAS: Shenmen(TF4), Endocrine(C018) Spleen(CO13), Stomach(CO4) Dachang(CO7), Sanjiao(CO17)	Acupuncture:30 min ear acupressure:1 day	1 session every day with a total of 10 sessions for 6 weeks, 1 week rest in-between 10 sessions.
	Acupuncture	Tianshu(ST25), Guanyuan(CV4) Zusanli(ST36), Fenglong(ST40) Sanyinjiao(SP6)	30 min	1 session every day with a total of 10 sessions for 6 weeks, 1 week rest in-between 10 sessions.
Shi et al. 2005, China	Warming acupuncture	Zhongwan(CV12), Shuifen(CV9) Qihai(CV6), Zhongji(CV3) Tianshu(ST25), Shuidao(ST28) Neiguan(PC6), Hegu(L14) Xuehai(SP10), Zusanli(ST36) Fenglong(ST40), Sanyinjiao(SP6)	40 min	l session every day with a total of 15 sessions for 4 weeks
	БА	Zhong wan(CV12/, Shuuen(CV9/) Qihai(CV6/, Zhongji(CV3) Tianshu(ST25), Shuidao(ST28) Neiguan(PC6), Hegu(L14) Xuehai(SP10), Zusanli(ST36) Fenglong(ST40), Sanyinjiao(SP6)	40 min	l session every day with a total of 15 sessions for 4 weeks
Yang et al. 2010, China	AR	Zhongwan(CV12), Tianshu(ST25) Guanyuan(CV4), Zusanli(ST36) Fenglong(ST40), Yinlingquan(SP9) Sanyinjiao(SP6), Pishu(BL20) Weishu(BL21), Ashixue	30 min	1 session daily with a total of 15 sessions for 7 weeks, 3 days rest between every session
Cabioglu et al. 2005, Turkey	EA	Body points: Hegu(LI 4), Tianshu(ST 25) Quchi(LI 11), Zusanli(ST 36) Neiting(ST 44	30 min	Body EA was performed everyday, and EA was performed every other day
Cabioglu et al. 2006, Turkey	EA	Body points: Quchi(LI II), Zusanli(ST 36) Neiting(ST 44)	30 min	Body EA application was performed daily for 20 days, and EA was applied to each ear on alternating days

TABLE 2: Continued.

Study ID (Country)	Style of acupuncture	Names of acupuncture points used	Retention time	Frequency & duration of Acupuncture sessions
Cabioglu et al. 2008, Turkey	EA	Body points: Hegu(LI 4), Quchi(LI 11) Tianshu(ST 25)Zusanli(ST 36) Taitong(Liv 3), Neiting(ST 44)	30 min	Body EA application was performed daily for 20 days, and EA was applied to each ear on alternating days
Darbandi et al. 2013, Iran	AR	Inrervention group: Tianshu(ST 25), Weidao(GB 28) Zhongwan(RN 12), Shuifen(RN 9) Guanyuan(RN 4), Sanyinjiao(SP 6) Excess group: Quchi(LI 11), Fenglong(ST 40) Deficiency group: Qihai(RN6), Yinlingquan(SP9)	20 min	Two treatmennt per week for a total of 6 weeks(12 treatments)
Fogarty et al. 2015, Australia	AR	Hegu(LI 4), Quchi(LI 11) Zusanli(ST 36), Neiting(ST 44) Taichong(LR 3) Auricular acupuncture: Shenmen(TF4)	NR	NR

LM: lifestyle modification; AAS: auricular acupoint stimulation; EA: electroacupuncture; ACE: acupoint catgut embedding; WA: warming acupuncture; AR: acupuncture and related therapies.

TABLE 3: Risk of bias assessment.

C 4-	Random sequence	11.5	Blinding of participants	Blinding of outcome	Incomplete outcome	Selective outcome
Study	generation	AHOCAHOH COHCEAHHEHE	and investigators	assessment	data addressed	reporting
Allison et al. 1995, USA	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Hsu et al. 2005, Taiwan	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Richards et al. 1998, Australia	Unclear risk	Low risk	High risk	Low risk	Low risk	Unclear risk
He et al. 2008, China	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Li et al. 2006, China	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Tong et al. 2011, China	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Hsu et al. 2009, Taiwan	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Hsieh et al. 2010, Taiwan	Unclear risk	Unclear risk	High risk	Low risk	High risk	Unclear risk
Hsieh et al. 2011, Taiwan	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Abdi et al 2012, Iran	Unclear risk	Unclear risk	High risk	Low risk	High risk	Unclear risk
Darbandi et al 2012, Iran	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
He et al 2012, China	Unclear risk	Unclear risk	High risk	Low risk	Unclear risk	Unclear risk
Lien et al 2012, Taiwan	Low risk	Low risk	High risk	Low risk	High risk	Unclear risk
Darbandi et al 2014, Iran	Low risk	Low risk	High risk	Low risk	Low risk	Unclear risk
Yeh et al. 2015, Taiwan	Low risk	Low risk	High risk	Low risk	High risk	Unclear risk
Chen et al. 2007, China	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Huang et al. 2011, China	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Tang et al. 2009, China	High risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Shi et al, 2006, China	Low risk	Unclear risk	High risk	Low risk	Unclear risk	Unclear risk
Hsu et al. 2005, Taiwan	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Güçel et al. 2012, Turkey	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Deng et al. 2014, China	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Hassan et al. 2014, Egypt	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
He et al. 2014, China	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Wang et al. 2013, China	High risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Sujung et al. 2014, South Korea	Low risk	Low risk	High risk	Low risk	Low risk	Unclear risk
Bu et al. 2007, China	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Shi et al. 2005, China	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Yang et al. 2010, China	High risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Cabioglu et al. 2005, Turkey	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Cabioglu et al. 2006, Turkey	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Cabioglu et al. 2008, Turkey	Unclear risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Darbandi et al. 2013, Iran	Low risk	Unclear risk	High risk	Low risk	Low risk	Unclear risk
Fogarty et al. 2015, Australia	Unclear risk	Low risk	High risk	Low risk	Low risk	Unclear risk

Commercian	Dairwing OR (050/ CI)	Number of mationts	Number of studies	Heterog	geneity test
Comparison	Pairwise OR (95% CI)	Number of patients	Number of studies	I ² (%)	p value
Body weight					
AR vs. LM	1.66(0.63 to 2.70)	496	10	55	0.02
AR vs. placebo	1.15(0.67 to 1.63)	833	14	65	0.0004
Combines therapies vs. PHA	2.44(-1.98 to 6.86)	80	1	-	-
Acupuncture vs. related therapies	0.25(0.00 to 0.49)	413	6	0	0.73
Combines therapies vs. acupuncture	1.56(0.07 to 3.05)	378	6	99	< 0.00001
BMI					
AR vs. LM	1.17(0.09 to 2.26)	314	6	74	0.002
AR vs. placebo	0.57(0.40 to 0.74)	830	12	63	0.002
Combines therapies vs. PHA	0.48(-0.90 to 1.86)	80	1	-	-
Acupuncture vs. related therapies	0.13(-0.06 to 0.32)	325	5	0	0.8
Combines therapies vs. acupuncture	0.77(-0.37 to 1.92)	158	4	88	< 0.00001

TABLE 4: Pairwise meta-analyses.

BMI: body mass index; LM: lifestyle modification; PHA: pharmacotherapy; AR: acupuncture and related therapies.

0.25tol.11), acupuncture (SMD: 1.28; 95% CrI, 0.43to2.06), combination of acupuncture and related theories (SMD: 1.44; 95% CrI, 0.64to2.38), and electroacupuncture (SMD: 0.60; 95% CrI, 0.03to1.22). Also, the combination of acupuncture and related theories and acupuncture alone were both superior to the acupuncture combined lifestyle modification in their ability to reduce body mass index (SMD = -1.76, 95% CrI = -2.96 to -0.62; SMD = -1.59, 95% CrI = -2.71to -0.34) (Table 5).

3.5. Ranking

3.5.1. Body Weight. Ranking of the different treatment methods was displayed Figure 3. The results suggested that, on the aspect of weight loss, combination of acupuncture and related therapies was ranked the optimal method, the best, (88.7%), followed by moxibustion with warming needle (87.8%), manual acupuncture (70.5%), acupoint catgut embedding (ACE,62.1%), auricular acupoint stimulation (AAS, 48.3%), electro acupuncture (EA,46.3%), pharmacotherapy (41.9%), acupuncture combined lifestyle modification (AR+LM,31.2%), placebo acupuncture/sham acupuncture (16.8%), and lifestyle modification (LM,6.4%) which was ranked as the worst.

3.5.2. BMI. The results suggested that, on the aspect of BMI, combination of acupuncture and related therapies was ranked the optimal method, the best, (90.2%), followed by manual acupuncture (83.3%), pharmacotherapy (64.7%), acupoint catgut embedding (58.6%), auricular acupoint stimulation (55.7%), electroacupuncture (52.1%), placebo acupuncture/sham acupuncture (25.1%), acupuncture combined lifestyle modification (16.9%), and lifestyle modification (3.4%) which was ranked as the worst.

3.6. Inconsistency Assessment

3.6.1. Body Weight. The Z test illustrates the inconsistency of the NMA specifically (Appendix 4). For the inconsistency test outcome of BW, 95% CI of 8 loops was included 0,

which reflected that no significant inconsistency was found. However, another 2 loops (ACE-Acupuncture-Combined therapies; ACE-Combined theories-EA) were found statistical inconsistency between direct and indirect comparisons.

3.6.2. BMI. For the inconsistency test outcome of body mass index, 95% CI of all loops (acupuncture -combined theories-EA; acupuncture-EA-placebo; AR+LM -EA- LM-placebo; ACE-acupuncture-EA; AAS-EA-LM; AAS-AR+LM-LM-placebo; AAS-EA-placebo) were included 0, which reflected that no significant inconsistency was found.

3.7. Safety. Ten RCTs [26–28, 31, 32, 35, 36, 38, 39, 45] reported adverse events, while no major complications were noticed in all included studies. Three included studies [35, 36, 39] reported that no adverse effects were noted in both experimental group and placebo group. In one included RCT, there were two patients reporting mild ecchymosis and one abdominal discomfort case reported as adverse events after electroacupuncture treatment; no case was reported in the lifestyle modification group [27]. In another study, there were seven subjects in group auricular acupoint stimulation and two subjects in group placebo had mild tenderness [32].

4. Discussion

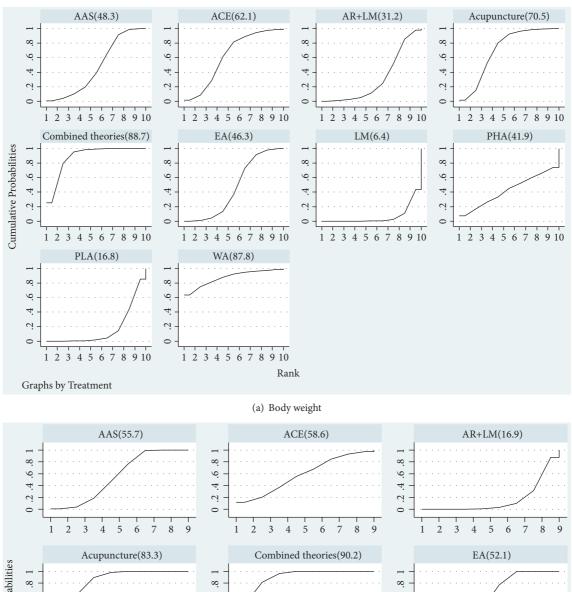
The aim of this study was to identify the efficacy and safety of acupoint stimulation therapy for obesity. In this NMA, the association of each acupuncture and related therapies with relative weight loss was compared by the combination of direct and indirect evidence from 34 RCTs in 2283 obese patients.

This study has three key findings. First, ranking graphs of the primary outcome suggested that the combination of acupuncture and related therapies was the most effective in losing weight and improving BMI. Second, compared with placebo or sham acupuncture, combination of acupuncture-related therapies, manual acupuncture, acupoint

TABLE 5: Results of network meta-analyses.

-1.11 (-4.01, 1.71) ACE							
0.72 (-0.93, 2.37) 1.82 (-1.08, 4.84)	4) AR+LM						
-1.45 (-4.11, 1.28) -0.36 (-1.82, 1.31)	31) -2.16 (-4.94, 0.61)	Acupuncture					
-2.40 (-5.16, 0.35) -1.28 (-2.96, 0.39)	39) -3.09 (-5.97, -0.33)	-0.92(-2.34, 0.30)	Combined theories				
-0.07 (-1.85, 1.74) 1.05 (-1.35, 3.48)	8) -0.77 (-2.61, 1.03)	1.38 (-0.77, 3.57)	2.33 (0.17, 4.56)	EA			
1.80 (0.21, 3.41) 2.90 (0.16, 5.72)	2) 1.09 (-0.27, 2.39)	3.26 (0.65, 5.88)	4.18(1.62, 6.83)	1.85 (0.37, 3.37)	ΓM		
0.06 (-5.43, 5.46) 1.11 (-3.79, 6.23)	3) -0.67 (-6.27, 4.70)	1.47 (-3.37, 6.35)	2.39 (-2.25, 7.16)	0.11 (-5.02, 5.13)	-1.76 (-7.18, 3.51)	Pharmacotherapy	
1.26 (0.46 , 2.06) 2.37 (-0.43, 5.18)	(8) 0.54 (-0.93, 2.03)	2.72 (0.06, 5.29)	3.65 (0.96, 6.34)	1.33 (-0.36, 2.96)	-0.55 (-2.02, 0.90)	1.20 (-4.13, 6.63)	Placebo
-3.47 (-8.46, 1.35) -2.31 (-7.69, 2.72)	72) -4.18 (-9.22, 0.67)	-1.99 (-7.32, 2.93)	-1.04 (-6.34, 3.96)	-3.40 (-8.10, 1.06)	-5.24 (-10.15, -0.55)	-3.50 (-10.48, 3.39)	-4.72 (-9.77, 0.07) WA
-0.08 (-1.54, 1.42) ACE							
0.96 (-0.08, 2.00) 1.03 (-0.58, 2.66)	(6) AR+LM						
-0.64 (-1.48, 0.35) -0.54 (-1.88, 0.84)	34) -1.59 (-2.71, -0.34)	Acupuncture					
-0.81 (-1.77, 0.12) -0.71 (-2.20, 0.70)	70) -1.76 (-2.95, -0.62)	-0.16(-1.01, 0.43)	Combined theories				
0.04 (-0.68, 0.78) 0.12 (-1.20, 1.45)	5) -0.92 (-1.88, 0.08)	0.67 (-0.11, 1.34)	$0.84\ (0.19, 1.58)$	EA			
1.31 (0.36, 2.30) 1.40 (-0.18, 2.96)	6) 0.34 (-0.36, 1.15)	1.94 (0.83, 3.00)	2.12 (1.07, 3.23)	1.28 (0.43, 2.11)	ΓM		
-0.31(-2.14, 1.55) $-0.21(-2.34, 1.87)$	37) -1.26 (-3.21, 0.79)	0.33 (-1.46, 2.01)	0.51(-1.06, 2.09)	-0.35 (-2.10, 1.39)	-1.62 (-3.54, 0.31)	Pharmacotherapy	
0.63 (0.25, 1.11) 0.72 (-0.71, 2.15)	5) -0.32 (-1.28, 0.69)	1.28(0.43, 2.05)	1.44 (0.64, 2.38)	0.60(0.03, 1.22)	-0.66(-1.58, 0.26)	0.95 (-0.85, 2.78)	Placebo

BMI: body mass index; LM: lifestyle modification; AAS: auricular acupoint stimulation; EA: electroacupuncture; ACE: acupoint catgut embedding; WA: warming acupuncture; AR: acupuncture and related therapies, combined therapies: combination of acupuncture and related therapies.



Cumulative Probabilities .4 .6 .8 8. 9. 9 4. 4. 7 7 7 0 0 0 5 5 7 5 3 4 8 3 6 8 6 PHA(64.7) PLA(25.1) LM(3.4) ∞. ∞ ∞ .4 .6 9 9. 4. 4. 7 7 3 2 5 4 Rank Graphs by Treatment (b) Body mass index

FIGURE 3: Surface under the cumulative ranking curves. LM: lifestyle modification; AAS: auricular acupoint stimulation; EA: electroacupuncture; ACE: acupoint catgut embedding; WA: warming acupuncture; AR: acupuncture and related therapies; combined therapies: combination of acupuncture and related therapies; PLA: placebo; PHA: pharmacotherapy.

catgut embedding, auricular acupuncture therapy, and electroacupuncture are all associated with higher odds of achieving weight loss. Third, combination of acupuncture and related therapies, manual acupuncture, pharmacotherapy, acupoint catgut embedding, auricular acupoint stimulation, and electroacupuncture were superior to lifestyle intervention.

Lifestyle modification, like diet intervention and physical activity, is recommended as safe and effective way to lose weight [60]. Results of direct and indirect evidence suggest acupuncture and related theories had significant beneficial effects in dealing with obesity compared with lifestyle modification. Both experimental and clinical data prove the efficacy of acupuncture for obesity [61]. Experimental data suggests that acupuncture exerts beneficial effects on weight loss [62, 63]. The majority of clinical evidence suggests that acupuncture and related therapies reduced more weight than sham control group [26, 28, 31, 32], which are consistent with our results. Previous animal studies have observed that the expression of obesity-related peptides was upregulated in the hypothalamus after acupuncture treatment, which induced less food intake and weight loss [62, 64, 65]. Similarly, significant decreases in plasma leptin level were observed after EA treatment in obese patients [46]. With regard to insulin level, several experimental studies reported that EA can improve insulin sensitivity [66, 67]. However, results from clinical trials regarding insulin levels are controversial. Cabioğlu MT reported that EA increased insulin level compared with control group [56], but Gucel F indicated that acupuncture decreased insulin level [46]. As to effects on lipid metabolism, acupuncture was reported to be effective in decreasing total cholesterol (TC), triglycerides (TG) and LDL-C concentrations [68, 69] of obese rat. Significant decreases in TC [55], TG [35], and LDL-C [55] were observed whereas no changes in HDL-C [55] levels were observed in clinical trials. Furthermore, experimental studies suggest that there was significant decrease in serum TNF α after EA [70]. Except for the noted mechanisms, EA can also induce white adipose tissue (WAT) browning via increasing uncoupling protein-1 (UCP1) gene expression [71].

This NMA has several attractive advantages. We focused on simple obesity patients without complication, which decreased the heterogeneity and improve the quality of this study. In addition, we compared acupuncture and acupuncture-related therapies with the first-line treatment for obesity-lifestyle modification with a Bayesian framework. The rank test of effectiveness provides data to favour acupuncture and acupuncture-related therapies. Lastly, we conducted a comprehensive search and included all eligible studies. We compared five different acupuncture treatments (manual acupuncture; electro acupuncture; auricular acupoint stimulation; acupoint catgut embedding; moxibustion with warming needle) in the clinical effectiveness in treating patients with obesity.

However, this study has several limitations. First, we failed to evaluate the safety of each acupoint stimulation therapy due to the limited data in primary studies. Future trials should report adverse events clearly to improve the quality of study design. Second, unaddressed concerns still exist regarding the long-term effects of using acupuncture

and acupuncture-related therapies on weight management in a clinical setting. The duration of acupuncture sessions and follow-up duration of most included trials ranging from four weeks to twelve weeks. Further clinical evaluation of acupuncture for obesity with longer follow-up appears warranted. Third, blinding of patients and researches was not applied among included studies and the included trials were mainly conducted in China, which may lead to publication bias [72]. Fourth, included study in our NMA lack of research compares the effectiveness between acupuncture, pharmacotherapy, and different types of combination of acupuncture. Further confirmatory comparative effectiveness trials should compare different types of combination of acupuncture. Except one study compared acupuncture and pharmacotherapy [29], additional research is needed to further explore. Finally, we use R-value to estimate the changes in standard deviations (SD), which might enlarge the SD compared with the originals values.

Overall, our results indicate that combination of acupuncture and related therapies ranks as the optimal method for reducing both weight and BMI. Further studies will clarify which combination of acupuncture and related therapies is better.

Data Availability

All data used to support the findings of this study are included within the supplementary information files.

Conflicts of Interest

All authors declare that they have no potential conflicts of interest.

Authors' Contributions

Yanji Zhang, Jia Li, and Guoyan Mo contributed equally to this work. Wei Huang, Zhongyu Zhou, and Yanji Zhang contributed to study design. Xianglin Chen, Hui Liu, and Teng Cai contributed to study selection. Xiangmin Tian, Teng Cai, Xian Zhang, and Xianglin Chen contributed to data collection and quality assessment. Figures 1–3 were prepared by Jia Li and Jing Liu. Tables 1–5 were prepared by Guoyan Mo. Appendices 1–4 were prepared by Jia Li. Huisheng Yang and Teng Cai were responsible for technical and language support. All authors have read and approved the final manuscript.

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Research Team Project (no. 2017ZXZ004), and Science and Technology Program of Hubei, China (no. 2016CFB221).

Supplementary Materials

Appendix 1: the PRISMA-NMA checklist treatment groups for included studies in a network meta-analysis. Appendix 2: search strategies for RCTs on acupuncture for obesity. Appendix 3: summaries of mean, standard difference, and sample size between treatment groups for included studies in a network meta-analysis. Appendix 4: inconsistency test. (Supplementary Materials)

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