

REVIEW

Acupuncture as an Adjunct to Lifestyle Interventions for Weight Loss in Simple Obesity: A Systematic Review and Meta-Analysis

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Objective: This study aims to evaluate the efficacy of acupuncture as an adjunct to lifestyle interventions on weight loss in simple

Methods: Potentially eligible studies that assessed anthropometric outcomes as well as glucose and lipid metabolism following acupuncture combined with lifestyle interventions in simple obesity patients were searched from eight databases from inception to October 2023. All analyses were performed using the Review Manager software v5.4 and Stata software v17.0.

Results: Herein, 25 randomized controlled trials (RCTs) involving 2018 patients were included. Generally, acupuncture combined with lifestyle interventions outperformed lifestyle interventions alone in terms of body weight (MD = -4.73, 95% CI [-6.32, -3.13], p < 0.00001), body mass index (BMI) (MD = -2.11, 95% CI [-2.75, -1.48], p < 0.00001), waist circumference (WC) (MD = -4.96, 95% CI [-6.89, -3.03], p < 0.00001, body fat percentage (BF %) (MD = -2.61, 95% CI [-4.51, -0.71], p = 0.007), fasting plasma glucose (FPG) (SMD = -0.66, 95% CI [-1.02, -0.31], p = 0.0002), fasting serum insulin (FINS) (SMD = -1.12, 95% CI [-1.44, -0.81], p < 0.00001), homeostasis model assessment-insulin resistance (HOMA-IR) (MD = -1.22, 95% CI [-1.50, -0.94], p < 0.00001), total cholesterol (TC) (SMD = -1.14, 95% CI [-1.83, -0.45], p = 0.001), triglyceride (TG) (SMD = -1.31, 95% CI [-2.07, -0.56], p = 0.0006), and low-density lipoprotein cholesterol (LDL-C) (SMD = -1.81, 95% CI [-3.11, -0.51], p = 0.006). Subgroup analysis based on three subgroup variables (types of acupuncture and lifestyle interventions as well as treatment durations) could partially explain the heterogeneity of the results.

Conclusion: Acupuncture demonstrated an enhancing effect on the benefits of lifestyle interventions for weight loss in patients with simple obesity, including reducing body weight and improving glucose and lipid metabolism. This finding suggests its potential value as a complementary therapy alongside lifestyle interventions in a clinical setting. However, further validation with rigorously designed and high-quality RCTs is still needed in the future.

Keywords: obesity, acupuncture, acupuncture therapy, meta-analysis, systematic review

Introduction

The global prevalence of obesity has risen dramatically, becoming a major public health concern. By 2023, estimates indicate that over 1 billion people worldwide were living with obesity. 1,2 The prevalence of obesity has also increased

among children and adolescents. In China, the prevalence of obesity is approximately 7.9% in adolescents and 16.4% in adults.³ According to the estimates by the World Obesity Atlas, the number of obese people aged over 5 years will achieve nearly 2 billion by 2035, accounting for 24% of the global population.⁴ As a result, obesity, a major cause of various metabolic diseases like hyperlipidemia, type 2 diabetes mellitus (T2DM), and cardiovascular disease (CVD),^{5–7} poses a significant challenge to global public health.

Simple obesity is defined as excessive accumulation of body fat due to intake of calorie exceeding the body's calorie expenditure, without significant endocrine, metabolic disorders, or other etiologies. It accounts for approximately 95% of all obesity cases worldwide.^{8,9} The negative effects caused by simple obesity are multifaceted. Besides weight gain resulting from increased size and number of adipocytes, simple obesity also leads to disorder in fat metabolism, increasing serum total cholesterol (TC), triglyceride (TG), and low-density lipoprotein cholesterol (LDL-C) as well as cardiovascular and cerebrovascular risks.^{10,11} In addition, simple obesity can result in insulin resistance (IR), augmenting blood glucose levels.¹² Moreover, simple obesity is a well-established independent risk factor for several cancers. It can significantly increase the risk of endometrial cancer, esophageal cancer, and colorectal cancer.¹³ Owing to its numerous negative consequences, it is imperative to explore effective and safe treatments for simple obesity.

Obesity is caused by the interaction of multiple factors. Among the recognized risk factors, unhealthy diets and sedentary lifestyles contribute to obesity most.³ As a result, lifestyle interventions such as dieting and exercise are the foundation of obesity management, which are proposed as the first-line treatment for obesity in Chinese, American, and European clinical guidelines.^{14–16} In addition to facilitating weight loss, adjusting unhealthy lifestyle habits through dieting and exercise also helps reduce the risk of other related diseases (such as T2DM, CVD, et al), highlighting their significance in obesity treatment and prevention.¹⁷ However, the curative effect of lifestyle interventions alone is gradual and moderate, at least six months of treatment is needed to achieve significant improvements.^{18,19} Therefore, good self-discipline and perseverance are critical factors to achieving weight loss when lifestyle interventions alone is used. Unfortunately, many people with obesity struggle to lose weight due to challenges with maintaining motivation and adherence to lifestyle changes. This highlights the importance of identifying safe and effective complementary therapies to support lifestyle interventions in obesity management.

Acupuncture, as a traditional Chinese medicine (TCM) therapy, has been widely used in treating many metabolic diseases such as T2DM, metabolic syndrome and polycystic ovary syndrome because of its effectiveness, safety, and economy. Preclinical studies in animals have shown promise for acupuncture's effectiveness in weight loss. These findings are echoed by numerous clinical trials suggesting acupuncture's benefit for weight management in patients with simple obesity, either when used alone (monotherapy) or as an adjunct to other therapies. Moreover, due to the critical involvement of obesity in the onset and progression of T2DM and cardiovascular accidents, studies also explored the effect of acupuncture in regulating abnormal glucose and lipid metabolism in obese patients. Enforcement effect of acupuncture on restoring blood glucose and lipid index to lifestyle interventions was found, that might be related to the neuro-endocrine-immune regulation by acupuncture. Besides, the safety and economy of acupuncture was also proved in previous studies, which is much safer and cheaper than the common anti-obesity drugs. Thus, it is worthwhile to consider incorporating acupuncture into obesity treatment procedures.

Obesity is a complex metabolic disease requiring a multi-pronged approach within management programs, with lifestyle interventions serving as the cornerstone. Acupuncture offers a safe, effective, and cost-effective therapeutic option in this context. Nonetheless, to our knowledge, no reviews have been published regarding the efficacy of acupuncture on weight loss in simple obesity, on the foundation of dieting or exercise. Thus, this study aims to evaluate and elucidate the efficacy of acupuncture as an adjunct to lifestyle interventions in reducing body weight and improving glucose and lipid metabolism in simple obesity. Our results are expected to provide reliable and evidence-based reference for the clinical application of acupuncture in obesity management programs.

Materials and Methods

Protocol and Registration

The protocol was registered on PROSPERO (No. CRD42023474447).

Search Strategy

This study was conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines (<u>Table S1</u>).³⁷ All eligible studies were searched from four English databases (PubMed, EMBASE, Web of Science and the Cochrane Library) and four Chinese databases (the Chinese Biomedical Literature Database, China National Knowledge Infrastructure, Wanfang, and the VIP Database) from inception to October 2023. Clinical condition (simple obesity and obesity), intervention methods (acupuncture, body acupuncture, manual acupuncture, and electro-acupuncture), and study type (randomized clinical trial) were used as the search terms in both the Chinese and English databases. <u>Table S2</u> outlines the complete search strategy.

Inclusion and Exclusion Criteria Study Type

All randomized controlled trials (RCTs) on acupuncture therapies combined with lifestyle interventions for simple obesity treatment were included. On the other hand, non-randomized clinical studies, animal experiments, conference papers, review papers, and case reports were excluded.

Types of Participants

Studies that involved participants diagnosed with simple obesity using any recognized diagnostic criteria were eligible for inclusion regardless of gender, age, and ethnic group.

Interventions and Comparisons

Studies with trial groups treated with acupuncture therapies [manual acupuncture (MA) or electro-acupuncture (EA)] combined with lifestyle interventions (dieting, exercise, or dieting plus exercise) regardless of the treatment duration and frequency, and control groups treated with lifestyle interventions alone or in combination with sham acupuncture (placebo) were included.

Outcomes

Primary outcomes were body weight and body mass index (BMI). Secondary outcomes included the following four indicators:

- 1. Anthropometric outcomes: waist circumference (WC) and body fat percentage (BF %).
- 2. Glucose metabolism: fasting plasma glucose (FPG), fasting serum insulin (FINS) and homeostasis model assessment-insulin resistance (HOMA-IR).
- 3. Lipid metabolism: TC, TG, high-density lipoprotein cholesterol (HDL-C) and LDL-C.
- 4. Adverse events (AEs): severe needle pain, bleeding, subcutaneous hematoma, fainting, and so on.

Study Selection and Data Extraction

Two researchers independently screened the literature. The studies failing to meet the inclusion criteria were excluded by evaluating the title, abstract, and full text. The studies finally included were determined through crosschecking, and a third-party reviewer were engaged in discussing and achieving consensus on differences. Two researchers independently extracted information from the literature into a predesigned form. The form contained general characteristics (first author, publication year, country, and sample size), characteristics of participants (age and BMI), traits of the intervention and control groups (dose of acupuncture, treatment duration, and follow-up time), and outcomes (Table S3).

Risk of Bias Assessment

Two researchers independently evaluated the risk of bias of the included studies using the Cochrane risk of bias tool.³⁸ The domains assessed included sequence generation, allocation concealment, blinding of participants, blinding of outcome assessors, incomplete outcome data, selective outcome reporting, and other sources of bias. The risk of bias in the literature was rated as "low", "high", and "uncertain". A third researcher assisted in the judgment in case of a disagreement.

Certainty Assessment

The quality of each piece of evidence was evaluated using the GRADE guideline (Grading of Recommendations Assessment, Development, and Evaluation).³⁹ Five factors that can reduce the quality of evidence include risk of bias, inconsistency, indirectness, imprecision, and publication bias. By evaluating these factors, the quality of evidence is divided into four levels including high, moderate, low, and very low.

Statistical Analysis

Statistical analysis was performed using the Review Manager software v5.4 and Stata software v17.0. Continuous data were summarized as mean differences (MD) or standardized mean difference (SMD) with 95% confidence intervals (CIs). The inter-study heterogeneity was assessed using I^2 statistics. The fixed-effect model was used when $p \ge 0.1$ and $I^2 \le 50\%$. Otherwise, the random-effect model was applied. In case of notable heterogeneity, sensitivity analysis was performed on all studies for detailed examination. Additionally, subgroup analysis was conducted to explore the potential effect modifiers (eg duration of intervention) and heterogeneity sources, that required at least 2 RCTs in each subgroup. Furthermore, publication bias was evaluated through funnel plot and Egger's test.

Results

Study Identification

The search strategy yielded 18336 studies. After importing all studies into Endnote X9 and removing duplicates (14737 remained), we screened titles and abstracts. This process yielded 1274 studies for full-text assessment, and ultimately, 25 studies were included in the meta-analysis (Figure 1).

Study Characteristics

Table 1 summarizes the characteristics of the included RCTs. The 25 RCTs involving 2018 patients were conducted in China, Malta, South Korea, Australia, Sudan and Iran. The sample sizes of these RCTs published between 2005 and 2023 range from 42 to 240 participants. All control groups received lifestyle interventions, such as dieting, exercise, or a combination of both. Moreover, some control groups received sham acupuncture. Besides, in addition to lifestyle interventions, the trial groups were treated with MA in 12 RCTs, ^{40−51} and with EA in the other studies. ^{52−64} Regarding the treatment duration, 18 RCTs and 7 RCTs lasted short-term ^{40−44,48,52−63} and long-term ^{45−47,49−51,64} (≤ 8 weeks and > 8 weeks), respectively. Except for 3 RCTs, ^{56,59,60} most of the included RCTs did not report follow-up data. Body weight, BMI, WC, BF %, FPG, FINS, HOMA-IR, TC, TG, HDL-C, LDL-C, and AEs were the outcomes considered in the meta-analysis.

Risk of Bias

Figure 2 depicts the risk of bias in all the included studies. Based on the applied quality criteria, methodological weaknesses were noted in most of the included RCTs. Regarding selection bias, 19 RCTs^{40,41,43–48,50,51,53–55,57,60,62–64} utilized random allocation but only 4 RCTs^{54,55,61,63} mentioned allocation concealment. The blinding of participants and personnel was not mentioned in most studies (except in 3 RCTs^{40,54,61}), indicating an unclear risk of bias. Similarly, only 3 RCTs reported blinding of outcome assessments. However, a low risk of bias was observed in the 'other bias' category for 22 RCTs. However, a low risk of bias was observed in the 'other bias' category for 22 RCTs.

Outcome Measurements

Body Weight

Seventeen studies recorded body weight of participants. $^{41,44,46,48-52,54-61,64}$ High heterogeneity was found among them ($I^2 = 80\%$, p < 0.00001). The random effects model revealed that acupuncture in combination with lifestyle interventions exerted a synergetic effect in decreasing body weight (MD = -4.73, 95% CI [-6.32, -3.13], p < 0.00001) (Figure 3).

Identification of studies via databases and registers

Records identified through databases Records removed before screening: searching (n = 18336) Duplicate records removed (n = 3599) Records excluded (n = 13463): Reviews (n = 669) Animal experiments (n = 1098) Conference summaries (n = 2301) Records for further screening Study ptorocols (n = 257) (n = 14737)Case reports (n = 342) Screening Meta-analysis (n = 480) Not relevant to the topic (n = 8316) Records excluded (n = 1249): Reports assessed for eligibility No lifestyle interventions (n = 30) (n = 1274)Unrelated outcomes (n = 21) Did not meet intervention or control criteria (n = 1192) Duplicate publication (n = 3)No full text (n = 3)Included Studies included in review (n = 25)

Figure I PRISMA flowchart for literature screening.

BMI

Twenty-two studies reported BMI. $^{40-51,54-61,63,64}$ A random-effect model was applied due to their significant heterogeneity ($I^2 = 93\%$, p < 0.00001). Compared to lifestyle interventions alone, acupuncture combined with lifestyle interventions exerted better effects in reducing BMI (MD = -2.11, 95% CI [-2.75, -1.48], p < 0.00001) (Figure 4).

WC

Fourteen studies compared the combination of acupuncture and lifestyle interventions (case group) to lifestyle interventions alone (control group) in obesity treatment using WC as an outcome. $^{42,44,46,48-52,55,56,58,61,63,64}$ The included studies had high heterogeneity ($I^2 = 90\%$, p < 0.00001). This meta-analysis found that acupuncture in conjunction with lifestyle interventions effectively reduced the WC (MD = -4.96, 95% CI [-6.89, -3.03], p < 0.00001; Figure 5A).

BF %

Nine studies reported BF % data. $^{42,45,47-49,52,56-58}$ A random-effect model was used ($I^2 = 96\%$, p < 0.00001). Compared to lifestyle interventions alone, acupuncture plus lifestyle interventions exerted a positive effect on BF % (MD = -2.61, 95% CI [-4.51, -0.71], p = 0.007) (Figure 5B).

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Table I Study Characteristics

Study	Country	Sample Size (n)	Age	(Year)	BMI (k	g / m²)	Treatment Regimen		Dose of Acupu	Treatment Duration	Follow- Up time	Outcomes	
		T/C	т	С	т	С	Т	С	ncture				
Tong et al, 2011 ⁴⁰	China	76/42	35.08±9.31	34.60±8.55	30.42±2.94	31.41±3.12	(1) + (4)	(6)	15 acupoints	5 weeks	NA	(b) (e) (h) (i) (l)	
Liang, 2013 ⁴¹	China	30/30	39.70 ± 3.59	37.50 ± 5.15	30.45 ± 3.25	28.32 ± 4.22	(1) + (4)	(4)	6 acupoints	22 days	NA	(a) (b) (f)	
Zhang and Wu, 2016 ⁴²	China	120/120	36.9 ± 9.86	38.8 ± 10.68	29.68 ± 0.56	29.80 ± 0.6	(1) + (3)	(3)	9 acupoints	8 weeks	NA	(b) (c) (d) (h) (i) (k) (l)	
Li et al, 2017 ⁴³	Susan	40/30	30.5±6.8	31.6±7.5	30.45±2.13	31.26±2.09	(1) + (5)	(5)	8 acupoints	4 weeks	NA	(b)	
Li et al, 2019 ⁴⁴	Australia	30/30	35.92±13.36	37.84±13.90	28.53±3.52	27.77±4.07	(1) + (5)	(5)	9 acupoints	6 weeks	NA	(a) (b) (c)	
Liu et al, 2021 ⁴⁵	China	46/47	43.9 ± 10.2	43.6 ± 8.8	30.46 ± 1.53	30.19 ± 1.36	(1) + (3)	(3)	7 acupoints	3 months	NA	(b) (d) (e) (f) (g) (h) (i)	
Min, 2021 ⁴⁶	South Korea	33/33	18–60	18–60	26.895±1.9369	27.093±4.0498	(1) + (3)	(3)	5 acupoints	16 weeks	NA	(a) (b) (c) (l)	
Zhang et al, 2021 ⁴⁷	China	30/30	39.13 ± 9.97	34.66 ± 9.87	31.63 ± 4.75	31.41 ± 2.43	(1) + (3)	(3)	3 acupoints	3 months	NA	(b) (d)	
Bi, 2022 ⁴⁸	China	35/35	36.71±10.18	36.43±9.81	29.29±1.10	29.87±1.64	(1) + (3)	(3)	15 acupoints	8 weeks	NA	(a) (b) (c) (d) (h) (i) (j) (k) (l)	
Qi and Shi, 2022 ⁴⁹	Malta	35/35	46.89±4.42	49.06±8.80	33.01±2.35	33.57±2.88	(1) + (3)	(3)	6 acupoints	12 weeks	NA	(a) (b) (c) (d) (h) (i) (k)	
Jia et al, 2023 ⁵⁰	China	40/40	37.2 ± 1.2	36.2 ± 3.2	28.73 ± 1.23	28.81 ± 1.18	(1) + (3)	(3)	8 acupoints	10 weeks	NA	(a) (b) (c) (h) (i) (j) (k)	
Yang et al, 2023 ⁵¹	China	47/47	34.37±7.66	33.47±8.18	30.58±4.91	29.26±4.59	(1) + (3)	(7)	16 acupoints	12 weeks	NA	(a) (b) (c)	
Li, 2005 ⁵²	China	35/33	38.97±10.88	40.09±11.37	NR	NR	(2) + (3)	(3)	8 acupoints	4 weeks	NA	(a) (c) (d) (h) (i) (j)	
Luo, 2007 ⁵³	China	30/30	17–55	20–54	27.9	27.5	(2) + (4)	(4)	10 acupoints	2 months	NA	(h) (i) (j) (k)	
Xing, 2009 ⁵⁴	China	31/30	28.16±7.17	28.37±7.13	27.83 ± 1.59	27.91 ± 1.47	(2) + (3)	(3)	10 acupoints	51 days	NA	(a) (b) (h) (i) (j) (k)	
Zhao and Shi, 2010 ⁵⁵	China	30/30	29.31 ± 8.28	30.30 ± 10.41	29.09 ± 3.64	27.84 ± 3.50	(2) + (3)	(3)	14 acupoints	8 weeks	NA	(a) (b) (c)	
Abdi et al, 2012 ⁵⁶	Iran	79/82	36.88±0.98	37.41±1.01	32.30±0.52	32.74±0.59	(1) + (4)	(6)	10 acupoints	6 weeks	6 weeks	(a) (b) (c) (d) (e) (h) (i) (j) (k)	
Wang and Xiao, 2014 ⁵⁷	China	45/45	48.8±3.5	49.4±3.9	30.1±2.2	29.8±2.0	(2) + (3)	(3)	12 acupoints	30 days	NA	(a) (b) (d) (l)	
Bae, 2016 ⁵⁸	China	30/30	32.97±8.032	29.33±7.485	29.363±3.222	30.417±4.564	(2) + (4)	(4)	19 acupoints	I month	NA	(a) (b) (c) (d)	
Xue et al, 2018 ⁵⁹	China	37/38	39.54 ± 6.96	36.47 ± 7.65	34.59 ± 6.96	33.61 ± 4.19	(2) + (3)	(3)	10 acupoints	8 weeks	3 months	(a) (b) (l)	
Yin et al, 2018 ⁶⁰	China	43/43	34.08±9.68	33.81±7.57	31.08±9.65	30.61±6.05	(2) + (3)	(3)	10 acupoints	8 weeks	3 months	(a) (b)	
Wang, 2020 ⁶¹	China	31/11	18–45	18–45	28.84 ± 2.67	28.75 ± 3.23	(2) + (4)	(4)	7 acupoints	5 weeks	NA	(a) (b) (c) (l)	
Zheng et al, 2021 ⁶²	China	30/30	35.65 ± 7.84	35.65 ± 7.81	28.00 ± 0.72	26.32 ± 0.39	(2) + (3)	(3)	12 acupoints	4 weeks	NA	(h) (i) (j) (k)	
Liang, 2022 ⁶³	China	30/30	37.00 ± 7.82	34.80 ± 8.35	28.00 ± 2.55	28.37 ± 3.08	(2) + (3)	(3)	13 acupoints	8 weeks	NA	(b) (c) (l)	
Ni et al, 2022 ⁶⁴	China	28/26	30.79 ± 8.8	30.50 ± 9.53	30.37 ± 4.75	31.12 ± 4.94	(2) + (3)	(3)	13 acupoints	12 weeks	NA	(a) (b) (c) (e) (f) (g)	

Notes: (1) Manual acupuncture; (2) Electro-acupuncture; (3) Dieting and exercise; (4) Dieting; (5) Exercise; (6) Sham acupuncture and dieting; (a) Body weight.

Abbreviations: C, control group; NA, not available; T, trial group; (b) BMI, body mass index; (c) WC, waist circumference; (d) BF %, body fat percentage; (e) FPG, fasting plasma glucose; (f) FINS, fasting serum insulin; (g) HOMA-IR, homeostasis model assessment-insulin resistance; (h) TC, total cholesterol; (i) TG, triglyceride; (j) HDL-C, high-density lipoprotein cholesterol; (k) LDL-C, low-density lipoprotein cholesterol; (l) Adverse events.

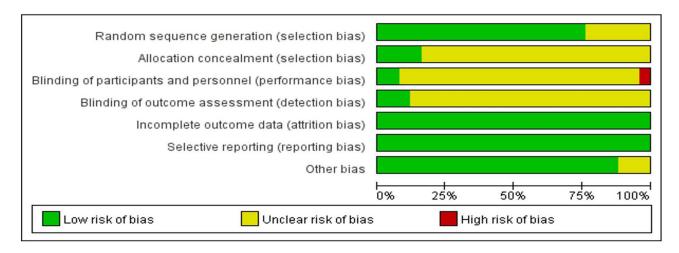


Figure 2 Risk of bias graph.

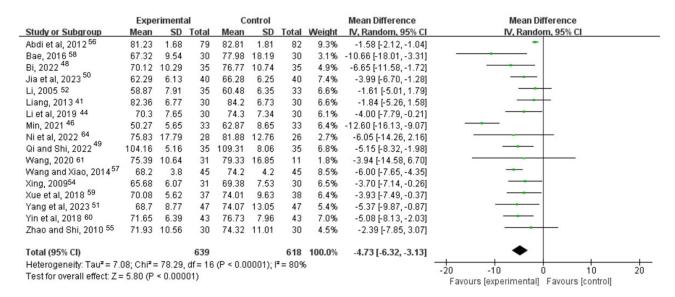


Figure 3 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on body weight.

FPG

Only four studies reported FPG data, 40,45,56,64 while high heterogeneity was found between them ($I^2 = 66\%$, p = 0.03). The results indicated a significant difference in FPG reduction between the trial and control groups (SMD = -0.66, 95% CI [-1.02, -0.31], p = 0.0002) (Figure 6).

FINS and HOMA-IR

Three studies 41,45,64 and two studies 45,64 compared the combination of acupuncture with lifestyle interventions against lifestyle interventions only in terms of changes in FINS and HOMA-IR, respectively. The fixed effects model was used due to the homogeneity of the results ($I^2 = 12\%$, p = 0.32; $I^2 = 6\%$, p = 0.30). Compared to lifestyle interventions alone, acupuncture, used as a supplementary method, significantly reduced FINS (SMD = -1.12, 95% CI [-1.44, -0.81], p < 0.00001) (Figure 7A) and HOMA-IR (MD = -1.22, 95% CI [-1.50, -0.94], p < 0.00001) (Figure 7B).

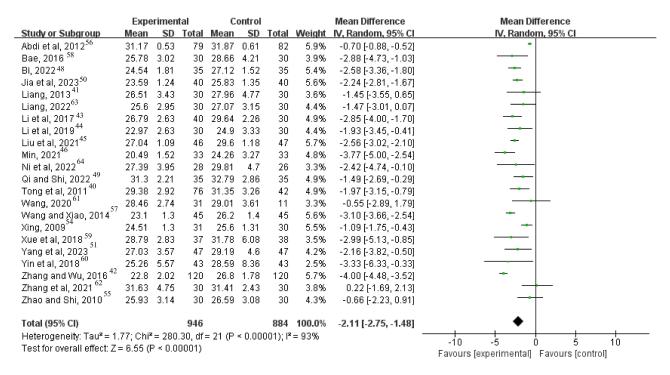
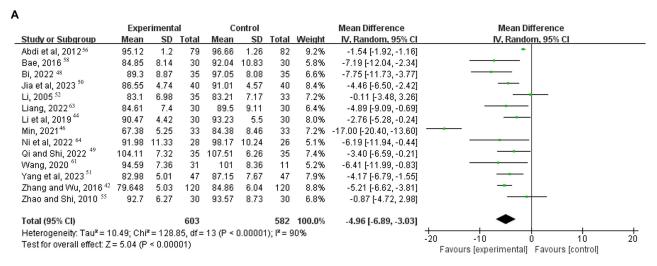


Figure 4 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on BMI. **Abbreviation**: BMI, body mass index.



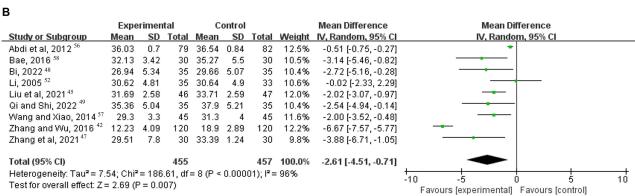


Figure 5 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on: (A) WC; (B) BF %. Abbreviations: BF %, body fat percentage; WC, waist circumference.

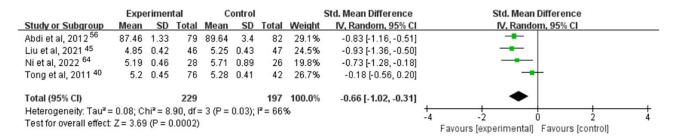


Figure 6 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on FPG. Abbreviation: FPG, fasting plasma glucose.

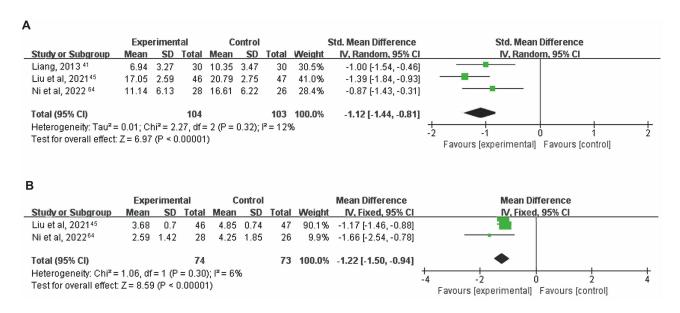


Figure 7 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on: (A) FINS; (B) HOMA-IR. Abbreviations: FINS, fasting serum insulin; HOMA-IR, homeostasis model assessment-insulin resistance.

TC and TG

Eleven studies reported TC and TG data. $^{40,42,45,48-50,52-54,56,62}$ High heterogeneity was detected in TC ($I^2 = 96\%$, p < 0.00001) and TG ($I^2 = 97\%$, p < 0.00001). According to the results, acupuncture in conjunction with lifestyle interventions is beneficial in reducing TC (SMD = -1.14, 95% CI [-1.83, -0.45], p = 0.001) (Figure 8) and TG (SMD = -1.31, 95% CI [-2.07, -0.56], p = 0.0006) (Figure 9).

HDL-C and LDL-C

Seven studies $^{48,50,52-54,56,62}$ and eight studies $^{42,48-50,53,54,56,62}$ reported changes in HDL-C and LDL-C, respectively. And significant heterogeneity was found in HDL-C ($I^2 = 90\%$, p < 0.00001) and LDL-C ($I^2 = 98\%$, p < 0.00001). The analysis confirmed that acupuncture plus lifestyle interventions was not superior to lifestyle interventions alone in increasing HDL-C (SMD = 0.47, 95% CI [-0.09, 1.03], p = 0.10) (Figure 10A), whereas reducing LDL-C (SMD = -1.81, 95% CI [-3.11, -0.51], p = 0.006) (Figure 10B).

Adverse Events

Only eight studies reported AEs. 40,42,46,48,57,59,61,63 Five studies reported no AEs. 40,46,48,57,59 Three studies reported mild AEs, 42,61,63 most of which were subcutaneous hematoma. All subjects fully recovered from AEs and completed the trial.

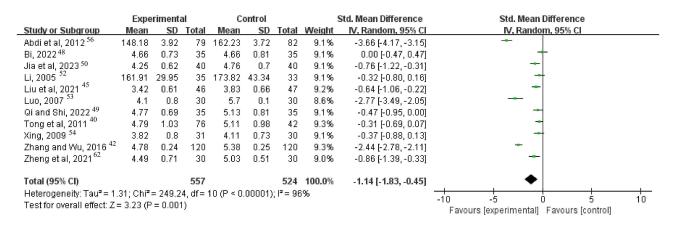


Figure 8 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on TC. **Abbreviation**: TC, total cholesterol.

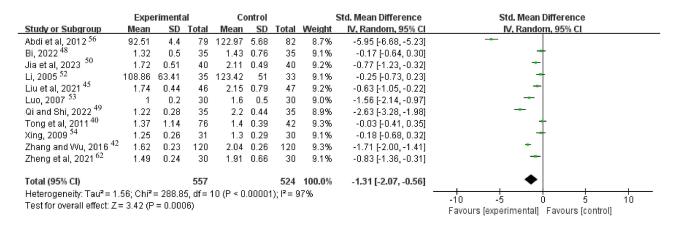
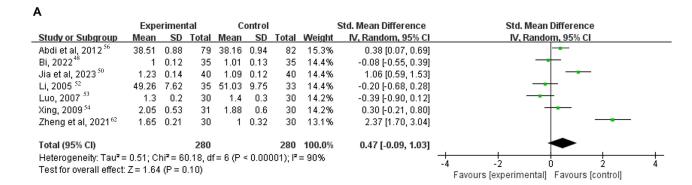


Figure 9 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on TG. **Abbreviation**: TG, triglyceride.

Subgroup Analysis

To further investigate the results, we conducted a subgroup analysis considering different types of acupuncture, lifestyle interventions, and treatment durations. In terms of acupuncture methods, both MA and EA, when combined with lifestyle interventions, exerted positive effects on body weight (MD = -5.60, 95% CI [-8.21, -2.99], p < 0.0001; MD = -3.99, 95% CI [-5.89, -2.09], p < 0.0001), BMI (MD = -2.41, 95% CI [-2.99, -1.83], p < 0.00001; MD = -1.80, 95% CI [-2.69, -0.92], p < 0.0001), and WC (MD = -6.23, 95% CI [-9.04, -3.42], p < 0.0001; MD = -3.05, 95% CI [-5.00, -1.10], p = 0.002), compared to lifestyle interventions alone. Regarding the effect on FPG, TC and LDL-C, EA combined with lifestyle interventions was superior to lifestyle interventions alone (SMD = -0.81, 95% CI [-1.09, -0.53], p < 0.00001; SMD = -1.59, 95% CI [-2.95, -0.23], p = 0.02; SMD = -1.60, 95% CI [-2.81, -0.39], p = 0.009). And MA in conjunction with lifestyle interventions is beneficial in reducing BF % and TG (MD = -3.62, 95% CI [-6.09, -1.15], p = 0.004; SMD = -0.97, 95% CI [-1.69, -0.26], p = 0.008) (Table 2 and Figures S1-S8).

Acupuncture showed a consistent additional effect on body weight (MD = -2.88, 95% CI [-5.07, -0.70], p = 0.010; MD = -5.22, 95% CI [-6.74, -3.69], p < 0.00001), BMI (MD = -1.75, 95% CI [-2.67, -0.84], p = 0.0002; MD = -2.30, 95% CI [-2.88, -1.72], p < 0.00001), and WC (MD = -3.38, 95% CI [-5.75, -1.01], p = 0.005; MD = -5.36, 95% CI [-7.84, -2.89], p < 0.0001) compared to other lifestyle approaches, including diet alone, exercise alone, or both combined. The difference was dieting or exercise alone showed no statistical significance in BF %, FPG, TC and TG while combination dieting and exercise showed no statistical significance in LDL-C (Table 3 and Figures S9-S16).



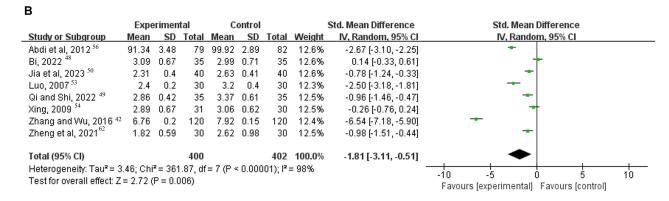


Figure 10 Forest plot of acupuncture combined with lifestyle interventions vs lifestyle interventions on: (A) HDL-C; (B) LDL-C. Abbreviations: HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol.

For treatment durations, there were significant reductions after short- and long-term acupuncture (≤ 8 weeks and ≥ 8 weeks, respectively) in body weight (MD = -3.88, 95% CI [-5.49, -2.27], p < 0.00001; MD = -6.66 95% CI [-10.12, -3.20], p = 0.0002), BMI (MD = -2.10, 95% CI [-2.96, -1.25], p < 0.00001; MD = -2.23, 95% CI [-2.86, -1.61], p < 0.00001), WC (MD = -3.72, 95% CI [-5.55, -1.89], p < 0.0001; MD = -7.00, 95% CI [-11.69, -2.31], p = 0.003), TC

Table 2 Subgroup Analysis of Different Acupuncture Interventions

Type of Acupuncture Interventions	Study (n)	Case (n)	MD/SMD 95% CI	Þ	l ² (%)	Model			
Manual acupuncture									
Body weight	7	500	-5.60 [-8.21, -2.99]	<0.0001	73	Random			
BMI	12	1081	-2.41 [-2.99, -1.83]	<0.00001	78	Random			
wc	7	680	-6.23 [-9.04, -3.42]	<0.0001	89	Random			
BF %	5	533	-3.62 [-6.09, -1.15]	0.004	92	Random			
FPG	2	211	-0.55 [-1.29, 0.19]	0.14	85	Random			
тс	6	671	-0.78 [-1.57, 0.01]	0.05	95	Random			
TG	6	671	-0.97 [-1.69, -0.26]	0.008	94	Random			
LDL-C	4	460	-2.03 [-4.55, 0.50]	0.12	99	Random			

(Continued)

Table 2 (Continued).

Type of Acupuncture Interventions	Study (n)	Case (n)	MD/SMD 95% CI	P	l ² (%)	Model			
Electro-acupuncture									
Body weight	10	757	-3.99 [-5.89, -2.09]	<0.0001	75	Random			
вмі	10	749	-1.80 [-2.69, -0.92]	<0.0001	88	Random			
WC	7	505	-3.05 [-5.00, -1.10]	0.002	56	Random			
BF %	4	379	-1.25 [-2.47, -0.02]	0.05	65	Random			
FPG	2	215	-0.81 [-1.09, -0.53]	<0.00001	0	Random			
тс	5	410	-1.59 [-2.95, -0.23]	0.02	97	Random			
TG	5	410	-1.74 [-3.48, -0.00]	0.05	98	Random			
LDL-C	4	342	-1.60 [-2.81, -0.39]	0.009	95	Random			

Notes: The random-effect model was used for the analysis of all indicators to make the results more reliable.

Abbreviations: CI, confidence interval; MD, mean difference; SMD, standard mean difference.

Table 3 Subgroup Analysis of Different Lifestyle Interventions

Type of Lifestyle Interventions	Study (n)	Case (n)	MD/SMD 95% CI	Þ	l ² (%)	Model			
Dieting or exercise alone									
Body weight	5	383	-2.88 [-5.07, -0.70]	0.010	47	Random			
вмі	7	571	-1.75 [-2.67, -0.84]	0.0002	76	Random			
WC	4	323	-3.38 [-5.75, -1.01]	0.005	66	Random			
BF %	2	221	-I.56 [-4.09, 0.96]	0.23	80	Random			
FPG	2	279	-0.52 [-1.16, 0.12]	0.11	85	Random			
тс	3	339	-2.24 [-4.52, 0.04]	0.05	98	Random			
TG	3	339	-2.50 [-5.67, 0.67]	0.12	99	Random			
LDL-C	2	221	-2.62 [-2.99, -2.26]	<0.00001	0	Random			
Combination of dieti	ng and exercis	e							
Body weight	12	874	-5.22 [-6.74, -3.69]	<0.00001	57	Random			
вмі	15	1259	-2.30 [-2.88, -1.72]	<0.00001	83	Random			
wc	10	862	-5.36 [-7.84, -2.89]	<0.0001	86	Random			
BF %	7	691	-2.89 [-4.94, -0.83]	0.006	91	Random			
FPG	2	147	-0.86 [-1.20, -0.52]	<0.00001	0	Random			
тс	8	742	-0.74 [-1.37, -0.11]	0.02	94	Random			
TG	8	742	-0.89 [-1.43, -0.34]	0.002	91	Random			
LDL-C	6	581	-1.55 [-3.21, 0.10]	0.07	98	Random			

Notes: The random-effect model was used for the analysis of all indicators to make the results more reliable.

(SMD = -1.34, 95% CI [-2.28, -0.39], p = 0.006; MD = -0.63, 95% CI [-0.89, -0.37], p < 0.00001), TG (SMD = -1.31, 95% CI [-2.31, -0.32], p = 0.010; SMD = -1.32, 95% CI [-2.39, -0.24], p = 0.02) and LDL-C (SMD = -2.13, 95% CI [-3.91, -0.35], p = 0.02; SMD = -0.87, 95% CI [-1.20, -0.53], p < 0.00001). Furthermore, after long-term acupuncture (>8 weeks), acupuncture assisted lifestyle interventions was superior to lifestyle interventions alone in reducing BF % (MD = -2.29, 95% CI [-3.20, -1.38], p < 0.00001) and FPG (SMD = -0.86, 95% CI [-1.20, -0.52], p < 0.00001) (Table 4 and Figures S17-S24). In the subgroup analysis, we found that a specific type of acupuncture, lifestyle intervention, or treatment duration may be the source of high heterogeneity.

Sensitivity Analysis

Regarding body weight and BMI, after any study was excluded, the range of change in MD are similar to the combined MD. Similarly, in terms of FPG, TC, and TG, the range of change in SMD is similar to that in the combined SMD. This indicated that each single study did not significantly affect the results, confirming the stability of the research findings. In spite of heterogeneity, the pooled result on these indicators was reliable. The results also showed that Min's study may be the source of the heterogeneity in WC. ⁴⁶ And the study by Zhang and Wu may have contributed to the heterogeneity in BF % and LDL-C levels. ⁴² When these two studies were excluded from the analysis, the range of change in SMD of the remaining studies are like the combined SMD in WC, BF % and LDL-C. Therefore, the interpretation of the results should be cautious. The details are in Tables S4-S10.

Table 4 Subgroup Analysis of Different Treatment Durations

Treatment Duration	Study (n)	Case (n)	MD/SMD 95% CI	Þ	l ² (%)	Model			
≤8 weeks									
Body weight	12	893	-3.88 [-5.49, -2.27]	<0.00001	72	Random			
BMI	15	1313	-2.10 [-2.96, -1.25]	<0.00001	94	Random			
WC	9	821	-3.72 [-5.55, -1.89]	<0.0001	82	Random			
BF %	6	689	-2,53 [-5.28, 0.21]	0.07	97	Random			
FPG	2	279	-0.52 [-1.16, 0.12]	0.11	85	Random			
тс	8	838	-1.34 [-2.28, -0.39]	0.006	97	Random			
TG	8	838	-1.31 [-2.31, -0.32]	0.010	97	Random			
LDL-C	6	652	-2.13 [-3.91, -0.35]	0.02	99	Random			
>8 weeks	>8 weeks								
Body weight	5	364	-6.66 [-10.12, -3.20]	0.0002	75	Random			
BMI	7	517	-2.23 [-2.86, -1.61]	<0.00001	60	Random			
wc	5	364	-7.00 [-11.69, -2.31]	0.003	91	Random			
BF %	3	223	-2.29 [-3.20, -1.38]	<0.00001	0	Random			
FPG	2	147	-0.86 [-1.20, -0.52]	<0.00001	0	Random			
тс	3	243	-0.63 [-0.89, -0.37]	<0.00001	0	Random			
TG	3	243	-1.32 [-2.39, -0.24]	0.02	93	Random			
LDL-C	2	150	-0.87 [-1.20, -0.53]	<0.00001	0	Random			

Notes: The random-effect model was used for the analysis of all indicators to make the results more reliable.

Publication Bias

More than 10 studies in each aspect of body weight, BMI, WC, TC, and TG were included. Funnel plot (<u>Figures S25-S29</u>) and Egger's test were used for publication bias assessment. The results indicated that it showed asymmetry in terms of body weight and WC in funnel plot. Furthermore, their results in Egger's test also suggested a significant publication bias between studies (p = 0.005 and p = 0.014, respectively). Despite the poor symmetry of the funnel plot, the results of Egger's test showed that no significant publication bias in BMI, TC and TG (p = 0.051; p = 0.967; p = 0.266).

Grading the Evidence

GRADE pro 3.6 software was used to assess the quality of evidence for all outcome indicators. The result indicated that the evidence for FPG, FINS and HOMA-IR were low quality, while the body weight, BMI, WC, BF %, TC, TG, HDL-C and LDL-C were very low quality (Table S11).

Discussion

Given its heavy burden on patients and society, obesity has become a global public health concern. Abnormal glucose and lipid metabolism are the main characteristic features of obesity that closely associated with some serious consequences like CVD. 65,66 Effective obesity management in obese patients goes beyond weight loss and must also address associated metabolic abnormalities due to the complex nature of the condition. Lifestyle interventions, therefore, serve as the foundation of a multi-component treatment approach. Building on prior research suggesting acupuncture benefits obese patients, 31–33 this study investigated its efficacy as an adjunct to lifestyle interventions for obesity treatment. We specifically focused on weight loss and its impact on glucose and lipid metabolism.

Summary of the Results

This meta-analysis is the first to evaluate the efficacy of acupuncture as an adjunct to lifestyle interventions for weight loss in simple obesity. In this study, we investigated the effectiveness of acupuncture when combined with lifestyle interventions, compared to lifestyle interventions alone. Results indicated that addition of acupuncture caused significant improvements in body weight, BMI, WC, BF %, FPG, FINS, HOMA-IR, TC, TG, and LDL-C. Importantly, acupuncture was associated with rare, mild, and self-resolving side effects.

Regarding weight loss, some studies reported that acupuncture might adjust appetite-regulating hormones, such as ghrelin and leptin, inducing a reduction in appetite among obese patients. Some appetite-related neurotransmitters, such as proopiomelanocortin, neuropeptide Y, agouti-related protein and the anorexigenic a-MSH peptide, were also found to be related to the effect of acupuncture on weight loss. Some appetite-related neurotransmitters, such as proopiomelanocortin, neuropeptide Y, agouti-related protein and the anorexigenic a-MSH peptide, were also found to be related to the effect of acupuncture on weight loss. Previous studies suggested that acupuncture may upregulate anti-inflammatory cytokines and downregulate pro-inflammatory cytokines. These abnormalities in cytokine regulation are thought to underlie the inflammatory mechanisms of obesity.

The benefits of acupuncture on glucose metabolism could be explained by the increase of whole-body glucose uptake induced by the activation of the parasympathetic and sympathetic nervous systems. Besides, IR, a key mechanism of glucose metabolism imbalance, is a common pathological state in obese patients, which could be reversed through acupuncture. Previous research also demonstrated the regulatory effect of acupuncture on lipid metabolism, including lipid decomposition, adipogenesis inhibition, and modulation of β - endorphins and leptin secretion. At 77–79

Despite our promising results, we observed high heterogeneity in some indicators, including body weight, BMI, WC, BF %, FPG, TC, TG and LDL-C. We conducted the subgroup analysis and sensitivity analysis to thoroughly investigate potential sources of heterogeneity. The results of subgroup analysis indicated that different types of acupuncture (MA and EA) and lifestyle interventions (dieting or exercise alone, and dieting plus exercise), and different treatment durations (\leq 8 weeks and > 8 weeks) could partially explain the heterogeneity between studies. Based on these results, clinicians can select long-term treatment (>8 weeks) of EA assisting dieting and exercise intervention to help patients with simple obesity lose weight and improve glucose and lipid metabolism. However, personalized treatment plans still need to be developed based on the physique and condition of patients in clinical practice.

The results of sensitivity analysis demonstrated that the pooled result had been proven to be reliable despite its heterogeneity in terms of body weight, BMI, FPG, TC, and TG. The variation likely stems from methodological flaws, such as differences in diagnostic criteria for simple obesity or variations in acupuncture parameters used across studies. However, it is important to note that these variations did not appear to influence the observed positive effect of acupuncture on these indicators. Besides, regarding of BF % and LDL-C, Zhang's study may be the cause for high heterogeneity. The diagnostic criteria and TCM syndrome types for simple obesity differed from other studies in this research. And the application of midnight-noon ebb-flow acupuncture method in Min's study is a specific operation method of acupoint selection according to syndrome differentiation and meridian to treat obesity, the which may be the source of heterogeneity for WC.

Applying the GRADE approach, the certainty of the evidence on the impact of acupuncture-assisted lifestyle interventions on weight loss in obese patients was very low. The quality of evidence for BMI and BF % was downgraded due to serious risks of bias and inconsistency, while for body weight and WC, publication bias further limited the certainty. Regarding glucose metabolism, the credibility of the evidence was low due to serious risks of bias, inconsistency, or imprecision. Additionally, the certainty of evidence for the impact on lipid metabolism was lowered to very low due to severe limitations from publication bias and inconsistency. Therefore, more high-quality RCTs are needed in this field to provide more solid evidence.

Our findings also found the common deficiencies of the RCTs of acupuncture on obesity in methodology. For instance, most of the included RCTs failed to report allocation concealment method, selective bias was unclear. Another potential limitation to consider is performance bias. Only four of the included RCTs mentioned implementing blinding procedures. This challenge is common in acupuncture trials, possibly due to the inherent nature of acupuncture therapy, which involves physical contact and specific needle manipulation techniques.

Comparison of Similar Studies

Consistent with our findings, multiple recent meta-analysis reported that acupuncture improved body weight, BMI, WC, and BF % in obese patients. 24–30 Notably, no meta-analysis examined the efficacy of acupuncture as an adjunct to lifestyle interventions on obesity. In other words, all studies overlooked the significance of lifestyle interventions as the first-line treatment for obesity. Fang's study compared the curative effects of acupuncture to placebo or no treatment groups on obesity, with or without lifestyle interventions. ABMI and adverse events are the observed outcomes in Fang's article. While in our study, other than the anthropometric outcomes including body weight and BMI, the results related to glucose and lipid metabolism are also analyzed. Zhong's study and Zhang's study examined the effects of acupuncture on weight loss in comparison to sham acupuncture. Gao's study established whether acupuncture combined with acupoint catgut embedding was superior to acupuncture alone in obesity treatment. Other three studies compared the effects of acupuncture treatment to non-acupuncture therapy on obesity. In contrast to previous research, we investigated the impact of acupuncture on both weight loss and its effects on glucose and lipid metabolism.

Unlike the above-mentioned studies, which overlooked the significance of lifestyle interventions as the first-line treatment for obesity, this study compared the impact of acupuncture combined with lifestyle interventions to that of lifestyle interventions alone in obesity treatment. In addition to the indicators directly related to obesity, the indicators related to glucose and lipid metabolism were also examined, providing comprehensive evidence on the effect of acupuncture on obesity. Performing subgroup analysis, sensitivity analysis, and publication bias to thoroughly explore potential sources of heterogeneity and conducting GRADE assessment for the evidence as well as analysis the clinical significance of the outcomes, are among the most important advantages of this research.

Implications for Clinical Practice

Aside from being a chronic metabolic disease, simple obesity is also an independent risk factor for T2DM and CVD. Obesity treatment not only aims to decrease weight, but also to help the body adapt to a normal status. Lifestyle interventions, proven to reduce body weight and significantly improve blood glucose and lipid metabolism, are the basis of obesity treatment. However, lifestyle interventions could only be independently effective after long-term maintenance, which could be challenging for most obese patients.

Acupuncture, as a non-drug therapy applied for thousands of years, has been widely utilized in the treatment of metabolic diseases. Herein, we demonstrated that acupuncture combined with lifestyle interventions had significant statistical difference on weight loss and glucose and lipid metabolism in obese patients. On the other hand, our findings confirmed that the effect size of acupuncture as an adjunct to lifestyle interventions in reducing body weight, BMI, WC, BF %, and HOMA-IR was higher than the minimal clinically important difference (MCID) for body weight (4.39 kg), 80 BMI (1.44 kg/m²), 81 WC (3.42 cm), 81 BF % (1.93%), 81 and HOMA-IR (0.05), 82 which is crucial for helping clinicians achieve their treatment goals. These evidence-based findings support the clinical application of acupuncture as an adjunct to lifestyle interventions. Other than this, acupuncture can help obese individuals partly increase their confidence and motivation to lose weight, thereby promoting their long-term adherence to a good lifestyle. By cultivating good lifestyle habits, the occurrence of obesity and its complications can be prevented or reduced.

Limitations of the Study

This study had some limitations. First, the sample sizes of the included studies were generally small except for one that included 240 obese patients. Second, most of the included studies had methodological deficiencies that might have affected the accuracy of our results. Third, due to the deficiency of available data in included studies, unfortunately we were unable to conduct a secondary analysis based on different TCM syndrome types of obesity. Fourth, most studies were published in Chinese and conducted in China, the generalization of these findings might be limited. Fifth, the original data of FPG, FINS, TC, TG, HDL-C and LDL-C used different units of measurement and the differences were resolved by SMD to represent the effect size. This limits the analysis of clinical significance of acupuncture as an adjunct to lifestyle interventions for obesity on these indicators. Finally, following the GRADE approach, the results of evidence quality were generally not optimistic, with all evidence quality being low or very low. Therefore, more well-designed RCTs are needed to further validate and draw more confident conclusions.

Implications for Future Research

Our meta-analysis demonstrates the positive effect of acupuncture as a supplementary therapy for lifestyle interventions on weight loss as well as glucose and lipid metabolism in obesity treatment. Regarding future clinical trials, researchers should adopt stricter trial designs to compensate for the quality deficiencies in methodology and evidence. In term of secondary analysis, researchers need to investigate the appropriate acupuncture therapy, including MA, EA and acupoint catgut embedding, for the management of obesity treatment via network meta-analysis. Furthermore, in addition to physiological improvements, it is also necessary to investigate acupuncture treatment in improving the psychological and social wellness of obese patients.

Conclusion

The findings of our study reveal that acupuncture, when used as an adjunct to lifestyle interventions, is effective in reducing body weight and improving glucose and lipid metabolism in simple obesity treatment. Moreover, the changes in body weight, BMI, WC, BF %, and HOMA-IR have clinical significance. However, considering the shortcomings of this study, such as the low quality of most included studies and the low or very low certainty of the evidence, future highquality and large-scale RCTs are needed to provide comprehensive and strong evidence for its clinical application in obesity management programs.

Data Sharing Statement

All relevant data are within the paper and its Supporting Information files.

Author Contributions

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

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Disclosure

The authors declare that they have no conflicts of interest in this work.

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