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Review

# Tai Ji Quan as antihypertensive lifestyle therapy: A systematic review and meta-analysis

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#### Abstract

*Background*: Professional health organizations are not currently recommending Tai Ji Quan alongside aerobic exercise to treat hypertension. We aimed to examine the efficacy of Tai Ji Quan as antihypertensive lifestyle therapy.

*Methods*: Tai Ji Quan interventions published in English and Chinese were included when they involved healthy adults, reported pre- and postintervention blood pressure (BP), and had a non-exercise/non-diet control group. We systematically searched 11 electronic databases for studies published through July 31, 2018, yielding 31 qualifying controlled trials. We (1) evaluated the risk of bias and methodological study quality, (2) performed meta-regression analyses following random-effects assumptions, and (3) generated additive models representing the largest possible clinically relevant BP reductions.

*Results*: Participants (n = 3223) were middle-aged (56.6 ± 15.1 years of age, mean ± SD) adults with prehypertension (systolic BP (SBP)=136.9 ± 15.2 mmHg, diastolic BP (DBP)=83.4 ± 8.7 mmHg). Tai Ji Quan was practiced 4.0 ± 1.4 sessions/week for 54.0 ± 10.6 min/session for 22.3 ± 20.2 weeks. Overall, Tai Ji Quan elicited significant reductions in SBP (-11.3 mmHg, 95%CI: -14.6 to -8.0;  $d_+ = -0.75$ ) and DBP (-4.8 mmHg, 95%CI: -6.4 to -3.1;  $d_+ = -0.53$ ) vs. control (p < 0.001). Controlling for publication bias among samples with hypertension, Tai Ji Quan trials published in English elicited SBP reductions of 10.4 mmHg and DBP reductions of 4.0 mmHg, which was half the magnitude of trials published in Chinese (SBP reductions of 18.6 mmHg and DBP reductions of 8.8 mmHg).

*Conclusion*: Our results indicate that Tai Ji Quan is a viable antihypertensive lifestyle therapy that produces clinically meaningful BP reductions (i.e., 10.4 mmHg and 4.0 mmHg of SBP and DBP reductions, respectively) among individuals with hypertension. Such magnitude of BP reductions can lower the incidence of cardiovascular disease by up to 40%.

Keywords: Blood pressure; Complementary medicine; Exercise training; Hypertension

# 1. Introduction

Hypertension is the most common cardiovascular disease risk factor, affecting nearly 50% of adults in the United States according to the 2017 American College of Cardiology/American Heart Association (ACC/AHA) Guidelines.<sup>1</sup> The ACC/AHA recommends aerobic exercise as antihypertensive lifestyle therapy

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\* Corresponding author. *E-mail address:* yin.2.wu@uconn.edu (Y. Wu). because it lowers blood pressure (BP) by  $5-8 \text{ mmHg.}^{1,2}$  Accordingly, adults with hypertension are encouraged to engage in moderate-to-vigorous intensity aerobic exercise on most, and preferably all, days of the week.<sup>1</sup> Unfortunately, most adults (72%) with hypertension do not adhere to these exercise recommendations for hypertension.<sup>3</sup>

Tai Ji Quan is a safe, low-impact, enjoyable, and inexpensive form of exercise accessible to individuals who for various reasons can not or choose not to engage in aerobic exercise.<sup>4-6</sup> Tai Ji Quan is a form of mind-body exercise that employs

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Cite this article: Wu Y, Johnson BT, Chen S, Chen Y, Livingston J, Pescatello LS. Tai Ji Quan as antihypertensive lifestyle therapy: a systematic review and metanalysis. J Sport Health Sci 2021;10:211–21. rhythmic physical motions while emphasizing relaxation and breathing techniques. Tai Ji Quan has numerous health benefits, with BP being one of the top health outcomes found in the literature.<sup>4,7,8</sup> In fact, Tai Ji Quan has been reported to elicit systolic BP (SBP) reductions as large as 17-19 mmHg and diastolic BP (DBP) reductions as large as 11-13 mmHg in primary-level Tai Ji Quan intervention studies published in English and Chinese.<sup>9-11</sup> Yet, due to limited evidence, professional committees and organizations such as the 2018 Physical Activity Guidelines Advisory Committee and ACC/AHA are not currently recommending Tai Ji Quan as antihypertensive lifestyle therapy alongside aerobic exercise, which is regarded as the standard of care.<sup>1,12,13</sup>

Three prior meta-analyses published in English have quantified the BP-lowering effects of Tai Ji Quan. Two of these meta-analyses reported clinically meaningful unstandardized mean effect sizes of -12.4 mmHg to -15.2 mmHg for SBP and -4.1 mmHg to -6.0 mmHg for DBP,<sup>14,15</sup> while the 3rd meta-analysis reported medium-to-large standardized mean effect sizes of -0.93 for SBP and -0.54 for DBP.<sup>16</sup> Despite these encouraging findings, these meta-analyses had the following limitations: (1) two of the 3 meta-analyses combined samples having various BP status, rendering it impossible to distinguish among adults who had normal BP from those who had hypertension;<sup>15,16</sup> (2) all meta-analyses had small sample sizes (k=9-12; n=536-832) and only included Tai Ji Quan intervention studies published in 2014 and before; and (3) all 3 meta-analyses did not disclose the statistical procedures used for the effect-size calculations as required by contemporary standards for systematic reviews and meta-analyses.<sup>17</sup> Furthermore, despite high heterogeneity ( $I^2$  ranged from 74% to 90%) for the 2 meta-analyses that gauged it<sup>15,16</sup>), moderator analyses were not performed to investigate if any study (e.g., trial location), sample (e.g., baseline BP status), or Tai Ji Quan intervention (e.g., frequency) characteristics influenced the BP response to Tai Ji Quan.<sup>14–16</sup>

To address the limitations in the literature, we conducted the largest meta-analysis to date to examine the efficacy of Tai Ji Quan as antihypertensive lifestyle therapy, while examining moderators of the BP response to Tai Ji Quan with meta-regression analysis, a contemporary statistical technique that can examine multiple moderators simultaneously.<sup>18–21</sup> In addition, we investigated which combination of study, sample, and Tai Ji Quan intervention characteristics generated the largest possible BP reductions with additive regression models to provide insights into the clinical utility of Tai Ji Quan as antihypertensive lifestyle therapy.

#### 2. Methods

#### 2.1. Selection criteria

The systematic review and meta-analytic procedures are registered at PROSPERO (registration number: CRD42019124262) and were conducted in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-analyses Statement.<sup>22</sup> According to 5 *a priori* inclusion criteria, trials were eligible if they: (1) enrolled adults  $\geq$ 18 years of age who were healthy other than their BP status, (2) explicitly stated that the intervention was Tai Ji Quan, (3) included a non-exercise/non-diet control group, (4) reported BP pre- and post-intervention for the Tai Ji Quan and control groups, and (5) were peer-reviewed articles published in either English- or Chinese-language journals. Trials were excluded if they (1) included samples with chronic disease (e.g., cardio-vascular disease, diabetes mellitus, cancer); (2) prescribed drug/supplements, dietary interventions, or another type of exercise in addition to Tai Ji Quan; or (3) involved a single Tai Ji Quan session (i.e., an acute intervention).

#### 2.2. Search strategy

Aided by a medical librarian (JL), we systematically searched 6 electronic databases in English and 5 electronic databases in Chinese from their inception until July 31, 2018 (Supplementary Table 1, Full search strategy, for each of the electronic databases queried). The team screened potential reports in duplicate by trained coders (YW, SC, and YC), first by title and abstract and then by full text. The team also searched for additional reports in reference lists of included trials, and reference lists of relevant reviews and meta-analyses.

#### 2.3. Data extraction and coded variables

Data were extracted using our standardized coding form and coder manual<sup>23,24</sup> adapted according to a previously published scale. This scale was specially designed to evaluate the frequency, intensity, and time principle of exercise prescription and instructional methods (e.g., breathing techniques, credentials of the instructors) of Tai Ji Quan interventions aimed at improving balance for older adults.<sup>25</sup>

We coded the study characteristics (e.g., trial location, publishing language), sample characteristics (e.g., baseline BP, body mass index), and Tai Ji Quan intervention characteristics (i.e., the frequency, intensity, and time exercise prescription and instructional methods). In addition, we coded BP measurement methods, such as BP measurement equipment and body position during measurement. Notably, we extracted information regarding the time between the last session of Tai Ji Quan and the post-intervention BP measurement, which may suggest the existence of the acute BP effects of the last session of exercise (i.e., postexercise hypotension<sup>21,26</sup>) and/or detraining since the BP benefits of exercise dissipate rapidly upon the cessation of training.<sup>27</sup> In our meta-analysis, we followed the Seventh Report of the Joint National Committee BP definitions of hypertension, prehypertension, and normal BP because the majority of research in this literature was conducted before release of the 2017 ACC/AHA Guidelines. Methodological study quality was assessed using an augmented version of the Downs and Black checklist (27 items) and was scored as the percentage of items satisfied out of a possible 32-point total.<sup>28</sup> Overall methodological study quality scores were grouped into low ( $\leq 16$  points, <50%), moderate (>16 to 25 points, 50%-79%), or high (>25 points,  $\geq 80\%$ ).<sup>29,30</sup> We used the Revised Cochrane Risk-of-Bias Tool for Randomized Trials

(RoB 2.0)<sup>31</sup> to assess risk of bias from domains including the randomization process, deviation from intended interventions, missing outcome data, measurement of outcomes, and selection of reported results for randomized controlled trials (RCTs). In addition, we used the Risk of Bias in Non-Randomized Studies of Interventions (ROBINS-I)<sup>32</sup> to assess risk of bias from domains, including the confounding, selection of participants, intervention classification, deviation from intended interventions, missing outcome data, measurement of outcomes, and selection of reported results for non-RCTs (NRCTs).

All data extraction and assessments of methodological study quality and risk of bias of included Tai Ji Quan interventions were conducted by 3 trained coders (YW, SC, and YC) independently and a high reliability was achieved (mean Cohen's  $\kappa = 0.87$  and Pearson's r = 0.88). All disagreements were resolved through discussion.

#### 2.4. Outcome and effect size calculations

The main outcome of the current meta-analysis is the BP response to Tai Ji Quan, defined as the difference between the change of BP in the Tai Ji Quan group and in the control group. The BP response to Tai Ji Quan was quantified as the standardized mean difference effect size (d) by calculating the mean difference in resting SBP or DBP between the Tai Ji Quan group and the control group post- vs. pre-intervention, divided by the pooled standard deviation (SD), correcting for small sample size bias and baseline differences between the groups.<sup>33,34</sup> When trials measured resting BP at multiple time points, we treated the BP value taken at the time point before and at the closest to the onset of the Tai Ji Quan intervention as the pre-intervention value, and the BP value taken at the time point after and closest to the end of the Tai Ji Quan intervention as the post-intervention value. Because we observed a large variation in the distribution of baseline SBP (the SD ranged from 3.3 to 25.3)<sup>35,36</sup> and DBP (the SD ranged from 3.6 to  $(17.1)^{11,37}$  across the included trials, we chose to use standardized effect sizes, which are more robust, less biased, and have better efficiency compared to unstandardized effect sizes across many statistical circumstances.<sup>38</sup> Negative d values indicate that Tai Ji Quan reduced BP more post- vs. pre-intervention compared to the control group. When reaching statistical significance (p < 0.05), d values were interpreted as insufficient (0 to -0.19), small (-0.20 to -0.49), medium (-0.50 to -0.79), and large ( $\leq$ -0.80) BP reductions.<sup>39</sup> Inconsistencies in d values were assessed with the Q statistic, which we transformed into the  $l^2$  statistic and its 95% confidence intervals (95%CIs).<sup>40,41</sup>  $I^2$  values range from low heterogeneity (0%) to high heterogeneity (100%).

# 2.5. Moderator analyses

We examined characteristics related to the study, sample, and Tai Ji Quan interventions as moderators of the BP response to Tai Ji Quan. Weighted regression models with maximum likelihood estimation of the random-effects weights (i.e., the inverse of the variance for each *d*) was used to explain variability in *d* values for SBP and DBP. In multiple-moderator models, we examined significant or trending moderators  $(p \le 0.10)$  from bivariate meta-regression analyses<sup>42</sup> in conjunction with the model coefficients and  $R^2$  values (i.e., between-study variance explained by a covariate) to determine the influence of individual moderators on the BP response to Tai Ji Quan.<sup>43</sup> The moving constant technique<sup>44</sup> estimated the magnitude of the weighted mean effect size (*d*<sub>+</sub>) and its 95%CI at different levels of interest for individual moderators while statistically controlling for the presence of other moderators held constant at their mean levels; the results are estimated, or predicted *d*<sub>+</sub> values, denoted as  $\hat{d}_{+}$ .<sup>44</sup>

For SBP and DBP, additive regression models were generated from the final multiple-moderator models that represented the greatest potential antihypertensive benefit from Tai Ji Quan. Individual moderators were assessed within the same model at the level that yielded the greatest BP reduction (i.e.,  $\hat{d}_+$  and 95%CI). To facilitate clinical interpretation for each moderator dimension and level of interest, we backconverted the standardized estimate (i.e.,  $\hat{d}_+$ ) into mmHg of BP change by multiplying the  $\hat{d}_+$  by the SD of baseline SBP (i.e., 15) and the SD of baseline DBP (i.e., 9) calculated on the combined sample from all included Tai Ji Quan trials.<sup>24</sup>

#### 2.6. Publication bias

We evaluated the potential for publication or other reporting biases in both SBP and DBP *d* values by (1) visually examining the distribution and asymmetry of funnel plots<sup>45</sup> and (2) using the tests of Begg and Mazumdar<sup>46</sup> and Egger et al.<sup>47</sup> tests, in addition to performing the Precision-Effect Test and Precision-Effect Estimate with standard error (SE) analysis to determine if potential publication bias needed to be controlled by including the SE of the BP response to Tai Ji Quan in the multiple-moderator models for SBP and DBP.<sup>48</sup>

#### 2.7. Sensitivity analyses

Notably, we did not restrict our sample to RCTs, and, in fact, there were more NRCTs (51.6%, k=16) than RCTs (48.4%, k = 15). Accordingly, we performed sensitivity analyses to compare the d values from RCTs and NRCTs.<sup>49</sup> There was no difference between the mean effect sizes of RCTs and NRCTs for SBP (p > 0.35) or DBP (p > 0.35), affirming the decision to combine RCTs and NRCTs for analyses. In addition, we paired study design with other variables included in the final multiple-moderator models and examined them as competing moderators of the BP response to Tai Ji Quan. Study design was the weaker, non-significant moderator when examined in tandem with the baseline SBP (p = 0.72), publishing language (p = 0.16), or the SE of the SBP response to Tai Ji Quan (p=0.31) in the SBP multiple-moderator model. It was also the weaker, non-significant moderator when examined in tandem with baseline DBP (p = 0.59) or publishing language (p = 0.42) in the DBP multiple-moderator model. These results reinforced our decision to combine RCTs and NRCTs in our final analyses.

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#### 2.8. Statistical computing

Analyses were performed using Stata Version 14.1 (Stata Corp., College Station, TX, USA) with macros for meta-analysis<sup>42,50</sup> and incorporated random-effects assumptions. Stata commands appear in Supplementary Table 2 (Stata commands). Descriptive statistics are reported as mean  $\pm$  SD unless stated otherwise. Two-sided significance level was p < 0.05.

# 3. Results

#### 3.1. Study search and study characteristics

We identified 31 controlled Tai Ji Quan trials that satisfied the inclusion criteria (Fig. 1). A list of included trials is available in Supplementary Table 3 (Reference list of included Tai Ji Quan trials).

The included Tai Ji Quan trials were published between 1997 and 2018 in English (k = 18, 58.1%) and Chinese (k = 13, 58.1%) 41.9%) language journals and each enrolled between 20 and 300 ( $n = 104 \pm 74$ , mean  $\pm$  SD) participants. Trials were RCTs (k = 15, 48.4%) or NRCTs (k = 16, 51.6%), and most examined BP as the primary outcome (k = 24, 77.4%). The attrition rate was reported or can be calculated for 18 of the Tai Ji Quan interventions (58.1%). On average, the attrition rate was  $16.8\% \pm 8.8\%$  (Supplementary Table 4, Study and Tai Ji Quan intervention characteristics). Among included Tai Ji Quan interventions, 24 interventions (77.4%) disclosed the content of the control groups (Supplementary Table 4), including routine daily activities (k = 14, 45.2%), usual care (k = 4, 12.9%), and attention control (k=6, 19.4%) such as educational lectures. The remaining 7 (22.6%) interventions did not disclose the content of the control groups. On average, the included Tai Ji Quan trials achieved "moderate"



Fig. 1. Flow chart detailing the systematic search of potential reports and selection process of included Tai Ji Quan trials.

methodological study quality on the augmented Downs and Black Checklist  $(50.1\% \pm 16.3\%)$ ,<sup>27</sup> despite widely varying scores (25.8% - 87.1%). The risk-of-bias ratings for individual trials appear in Supplementary Table 5 (Risk of bias of RCTs) and Supplementary Table 6 (Risk of bias of NRCTs), respectively. The majority of the RCTs (k = 12, 80%) exhibited high levels of risk of bias based on RoB 2.0.<sup>31</sup> In particular, deviation from intended interventions was the domain with the highest percentage of trials that were rated as high risk of bias (k = 11, 73.3%). In parallel, the majority of the NRCTs (k = 11, 73.3%)68.8%) exhibited serious risk of bias based on ROBINS-I.<sup>32</sup> In particular, confounding was the domain with the highest percentage of trials that were rated as serious risk of bias (k = 10, k = 10)62.5%); meanwhile, deviation from intended interventions was the domain with the highest percentage of trials that did not provide enough information for judgments to be made (k = 11, 68.8%).

#### 3.2. Sample characteristics

Baseline sample characteristics were similar between the Tai Ji Quan (n = 1654) and control (n = 1569) groups (p > 0.05)(Supplementary Table 7, Baseline sample characteristics). On average, participants (n = 3223) were middle-aged (56.6  $\pm$  15.1 years) adults with prehypertension (SBP =  $136.9 \pm 15.2 \text{ mmHg}$ /  $DBP = 83.4 \pm 8.7 \text{ mmHg}$ ) and were predominantly women  $(69.5\% \pm 21.6\%)$ . Among the included samples, based on baseline BP values and use of BP medications.<sup>51</sup> 16 trials (51.6%) involved samples (n = 1750) with hypertension (SBP = 148.4  $\pm$ 8.9 mmHg/DBP = 88.3  $\pm$  8.3 mmHg). Meanwhile, 10 trials (32.3%) involved samples (n=854) with prehypertension  $(SBP = 130.4 \pm 5.6 \text{ mmHg/DBP} = 80.9 \pm 3.6 \text{ mmHg})$ , and 5 trials (16.1%) involved samples (n=619) with normal BP  $(SBP = 113.2 \pm 5.4 \text{ mmHg/DBP} = 72.5 \pm 3.9 \text{ mmHg})$ . A total of 8 trials (25.8%) provided information regarding medication use. Of these, 5 trials reported that no participants were taking BP medications, while all participants had hypertension.<sup>37,52–5</sup> Just 2 trials disclosed the percentage of participants who were taking specific categories of BP medications, with calcium channel blockers being the most popular (i.e., 52%-55%).<sup>56,57</sup> In addition, 1 trial reported that all of its participants had dyslipidemia and were taking statins.<sup>58</sup>

#### 3.3. Tai Ji Quan intervention characteristics

Table 1 provides the details of the Tai Ji Quan intervention characteristics. Tai Ji Quan was practiced, on average,  $4.0 \pm$ 1.4 sessions/week for  $54.0 \pm 10.6$  min/session for  $22.3 \pm 20.2$ weeks. Tai Ji Quan interventions included, on average,  $30.4 \pm$ 25.4 forms following Yang (k=15, 48.4%), Chen (k=1, 3.2%),<sup>52</sup> or undisclosed (k=15, 48.4%) styles; however, less than one-third of the trials disclosed the names of the Tai Ji Quan forms (k=8, 25.8%) or adopted progression methods (Table 1) to facilitate the learning of Tai Ji Quan practice (k=10, 32.3%). Few trials reported emphasis on the 3 fundamental elements of Tai Ji Quan practice, including movement principles (k=4, 12.9%), breathing techniques (k=6, 19.4%),

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Tai Ji Quan intervention characteristics.

Methods	k	Mean $\pm$ SD (range)	Reporting rate (%) <sup>a</sup>
Ex R <sub>x</sub> methods items			
Frequency	31	$4.0 \pm 1.4 (2.0 - 7.0)$	100.0
(sessions/week)			
Intensity <sup>b</sup>	12	_	38.7
Disclosed	1	_	
Monitored, not disclosed	11	_	
Time (min/session)	29	$54.0 \pm 10.6  (30.0 - 65.0)$	93.5
Length of intervention	31	$22.3 \pm 20.2 (6.0 - 104.0)$	100.0
(weeks)			
Instructional methods items			
Style <sup>b</sup>	16	_	51.6
Yang style	15	_	
Chen style	1	_	
Number of forms	24	$30.4 \pm 25.4  (6.0 - 108.0)$	77.4
Form names	8	_	25.8
Movement principles	4	_	12.9
emphasized			
Breathing techniques	6	_	19.4
emphasized			
Relaxation emphasized	3	_	9.7
Progression <sup>b</sup>	10	_	32.3
Designated leaning phase	8	_	
Gradually increased	2	_	
duration			
Number of instructors <sup>b</sup>	14	$1.4 \pm 0.9 \ (1.0 - 4.0)$	45.2
Single instructor	10	_	
Multiple instructors	4	—	
Credentials of	15	—	48.4
<i>instructors</i> <sup>b</sup>			
Experienced	4	—	
Master or expert	3	—	
Professional	5	—	
Trained, qualified,	6	—	
or certified			
Unsupervised practice,	5	—	16.1
encouraged, but did not			
disclose actual time spent <sup>b</sup>			

Notes: Summary statistics are based on 31 Tai Ji Quan trials (k) and are presented as mean  $\pm$  SD, unless otherwise stated; Range represents the minimum, maximum values reported for the particular item. — indicates that an item is not applicable.

<sup>a</sup> Reporting rate is expressed as a percentage of trials that reported the specific item out of the total number of trials (k = 31).

<sup>b</sup> For these Tai Ji Quan intervention characteristics, the subcategories are listed with the frequency they appeared among those trials that reported the item. Authors used 1 or multiple keyword(s) to describe the credentials of the instructors.

Abbreviation: Ex  $R_x$  = exercise prescription.

or relaxation (k = 3, 9.7%). Only 1 trial (3.2%) reported that all 3 elements were emphasized.<sup>37</sup>

Among included Tai Ji Quan trials, only 1 trial (3.2%) disclosed the average heart rate (HR) during Tai Ji Quan practice (i.e., 63.7% of the age-predicted maximum HR.<sup>59</sup> Meanwhile, 10 trials (32.3%) monitored and instructed participants to exercise at various HR ranges that ranged from 40% to 85% of the age-predicted maximum HR, or between 110 beats/min and 130 beats/min, but they did not disclose the actual HR values. Only 1 trial stated that HR was recorded during Tai Ji Quan exercise but provided no further information.<sup>60</sup> All trials (k = 31, 100.0%) included supervised Tai Ji Quan exercise; yet, only about one-half of the trials disclosed the number (k = 14, 45.2%) or the credentials (k = 18, 58.1%) of the instructors (Table 1). In addition, 5 trials (k = 5, 16.1%) recommended unsupervised home practice without disclosing the actual time spent practicing. Last, in our sample, 2 trials (6.5%) specified that there were no adverse events; 1 trial (3.2%) indicated that no subjects withdrew from the study due to injuries from Tai Ji Quan practice. The rest of the trials (k = 28, 90.3%) did not provide any information regarding adverse events (Supplementary Table 4).

#### 3.4. Resting BP assessment

Most trials (k=21, 67.7%) reported the BP measurement instrument used. These included automated/digital units (k=10, 32.3%) or manual sphygmomanometers (k=11, 35.5%). About one-third of the trials (k=10, 32.3%) reported the body position during the BP measurements, which included the seated (k=9, 29.0%) or supine (k=1, 3.2%) position. Yet, 11 trials (35.5%) did not report any details of the resting BP measurement procedures. In addition, none of the Tai Ji Quan trials specified the time between the post-intervention BP measurements and the last Tai Ji Quan session.

# 3.5. Tai Ji Quan as stand-alone antihypertensive therapy and potential mechanisms

Overall, Tai Ji Quan elicited moderate to large reductions in SBP (-11.3 mmHg, 95%CI: -14.6 to -8.0;  $d_+ = -0.75$ ) and DBP (-4.8 mmHg, 95%CI: -6.4 to -3.1;  $d_+ = -0.53$ ) compared to controls (p < 0.001) (Supplementary Fig. 1, Funnel plots of the SBP and DBP response to Tai Ji Quan *vs.* control; Supplementary Fig. 2, Forest plots of the SBP and DBP response to Tai Ji Quan *vs.* control). Of note, the standardized effect sizes for SBP ( $I^2 = 88.3\%$ , 95%CI: 84.6%-91.2%) and DBP ( $I^2 = 83.9\%$ , 95%CI: 78.1%-88.2%) lacked homogeneity. In addition, the mean effect sizes of RCTs and NRCTs

Table 2

Moderator and additive models: SBP response to Tai Ji Quan  $(k=31)^a$ .

were similar for SBP (-0.72 vs. -0.78, p = 0.83) and DBP (-0.45 vs. -0.60, p = 0.41), which supported our decision to combine RCTs with NRTCs in our final analyses.

In relation to the mechanisms of the BP-lowering effects of Tai Ji Quan (Supplementary Table 4), 15 trials (48.4%) discussed potential mechanisms of the BP-lowering effects of Tai Ji Quan. However, only 5 trials (16.1%) measured one or more of the mechanisms discussed or proposed. In addition, only 1 trial (3.2%) examined the relationship between the change of BP in response to Tai Ji Quan and the measured mechanism. Specifically, Pan and colleagues<sup>61</sup> found that the change of plasma NO, CO, and H<sub>2</sub>S levels were negatively correlated with the change of SBP and mean arterial pressure.

### 3.6. Publication bias

The funnel plot (Supplementary Fig. 1), test of Begg, and test of Egger all suggested that there was significant publication or other reporting bias for the SBP response to Tai Ji Quan (test of Begg: z = -2.97, p = 0.003; test of Egger: t = -3.91, p = 0.001).<sup>45–47</sup> Meanwhile, both the funnel plot and the test of Egger suggested that there was significant publication or other reporting bias for the DBP response to Tai Ji Quan (t = -2.37, p = 0.03), but the test of Begg did not (z = -1.41, p = 0.16).<sup>45–47</sup> Following the Precision-Effect Test and Precision-Effect Estimate with SE<sup>48</sup> analysis, potential publication bias was apparent (p = 0.02) only for the SBP response to Tai Ji Quan; thus, publication basis was controlled for in the multiple-moderator model for SBP.

#### 3.7. Multiple-moderator models

When controlling for publication bias, SBP reductions were greater among Tai Ji Quan trials published in Chinese (16.7 mmHg, 95%CI: 12.5–20.9) than those published in English (8.4 mmHg, 95%CI: 4.8–11.9; p = 0.005) (Table 2). In addition, SBP was reduced more in samples with the highest baseline SBP (p = 0.04): 14.4 mmHg (95%CI: 11.3–17.7) for samples with hypertension, 11.3 mmHg (95%CI: 8.3–14.1)

Moderator dimension/level	<i>d</i> <sub>+</sub> (95%CI)	β	p	$\Delta$ SBP (mmHg, 95%CI)
Publishing language				
Published in Chinese $(k=13)$	-1.11 (-1.39  to  -0.83)	-0.391	0.005	-16.7 (-20.9 to -12.5)
Published in English $(k = 18)$	-0.56 (-0.79 to -0.32)			-8.4 (-11.9 to -4.8)
Baseline SBP of sample (mmHg)		-0.258	0.045	
Normal: $113 \pm 5 \ (k=5)$	-0.55 (-0.88 to -0.22)			-8.3 (-13.2 to -3.3)
Prehypertension: $130 \pm 6 \ (k=10)$	-0.75 (-0.94 to -0.55)			-11.3 (-14.1 to -8.3)
Hypertension: $148 \pm 9 \ (k=16)$	-0.96 (-1.18 to -0.75)			-14.4 (-17.7 to -11.3)
Publication bias (SE of the SBP response to Tai Ji Quan)			0.017	
Additive regression model: (1) among samples with				
hypertension and (2) controlling for publication bias				
Published in Chinese	-1.24 (-1.54 to -0.94)			-18.6 (-23.1 to -14.1)
Published in English	-0.69(-0.96  to  -0.41)			-10.4 (-14.4 to -6.2)

Note: Baseline SBP is presented as mean  $\pm$  SD.

<sup>a</sup> Multiple  $R^2$  (variance explained by model, adjusted for number of moderators) = 45.3%.

Abbreviations:  $\Delta$  = change;  $\beta$  = standardized coefficient represents unique variance explained by moderator; CI = confidence interval;  $\hat{d}_+$  = standardized estimate; SBP = systolic blood pressure; SE = standard error.

#### Antihypertensive effect of Tai Ji Quan

Table 3 Moderator and additive models: DBP response to Tai Ji Quan  $(k=31)^{a}$ .

Moderator dimension/level	$\hat{d}_+$ (95%CI)	β	р	Δ DBP (mmHg, 95%CI)
Publishing language				
Published in Chinese $(k = 13)$	-0.85 (-1.04 to -0.67)	-0.495	< 0.001	-7.7 (-9.4  to  -6.0)
Published in English $(k=18)$	-0.32 (-0.46 to -0.18)			-2.9(-4.1  to  -1.6)
Baseline DBP of sample (mmHg)		-0.464	< 0.001	
Normal: $73 \pm 4 \ (k=5)$	-0.31 (-0.49 to -0.13)			-2.8 (-4.4 to -1.2)
Prehypertension: $81 \pm 4$ ( $k = 10$ )	-0.53 (-0.64 to -0.41)			-4.8 (-5.8  to  -3.7)
Hypertension: $88 \pm 8 \ (k = 16)$	-0.71 ( $-0.83$ to $-0.59$ )			-6.4 (-7.5 to -5.3)
Additive Model: (1) among samples with hypertension				
Published in Chinese	-0.98 (-1.15 to -0.80)			-8.8 (-10.4 to -7.2)
Published in English	-0.44 (-0.61 to -0.28)			-4.0 (-5.5 to -2.5)

Note: Baseline DBP is presented as mean  $\pm$  SD.

<sup>a</sup> Multiple  $R^2$  (variance explained by model, adjusted for number of moderators) = 65.9%.

Abbreviations:  $\Delta$  = change;  $\beta$  = standardized coefficient represents unique variance explained by moderator; CI = confidence interval;  $\hat{d}_+$  = standardized estimate; DBP = diastolic blood pressure.

for samples with prehypertension, and 8.3 mmHg (95%CI: 3.3-13.2) for samples with normal BP. Collectively, these moderators accounted for 45.3% of the variance in the SBP response to Tai Ji Quan. The additive regression model revealed that, when controlling for publication bias among samples with hypertension, Tai Ji Quan trials published in Chinese elicited SBP reductions of 18.6 mmHg (95%CI: 14.1-23.1), nearly twice the magnitude of the SBP reductions of 10.4 mmHg (95%CI: 6.2-14.4) for Tai Ji Quan trials published in English (Table 2).

DBP reductions were greater among Tai Ji Quan trials published in Chinese (-7.7 mmHg, 95%CI: -9.4 to -6.0) than those published in English (-2.9 mmHg, 95%CI: -4.1 to -1.6; p < 0.001) (Table 3). In addition, DBP was reduced more in samples with the highest baseline DBP (p < 0.001): 6.4 mmHg (95%CI: 5.3-7.5) for samples with hypertension, 4.8 mmHg (95%CI: 3.7-5.8) for samples with prehypertension, and 2.8 mmHg (95%CI: 1.2-4.4) for samples with normal BP. Collectively, these moderators accounted for 65.9% of the variance in the DBP response to Tai Ji Quan. The additive regression model revealed that, among samples with hypertension, Tai Ji Quan interventions published in Chinese elicited DBP reduction of 8.8 mmHg (95%CI: 7.2-10.4), more than twice the magnitude of the DBP reductions of 4.0 mmHg (95%CI: 2.5-5.5) for Tai Ji Quan trials published in English (Table 3).

# 4. Discussion

Our meta-analysis examined the efficacy of Tai Ji Quan as antihypertensive lifestyle therapy with the largest sample size examined to date. We found that Tai Ji Quan practiced ~4 sessions/week for ~60 min/session for ~22 weeks, on average, produced BP reductions of 5-11 mmHg. Our novel finding, after controlling for publication bias among samples with hypertension, was that Tai Ji Quan trials published in English elicited SBP reductions of 10 mmHg and DBP reductions of 4 mmHg; for those published in Chinese, Tai Ji Quan trials elicited SBP reductions of 19 mmHg and DBP reductions of 10 mmHg. Even the more conservative magnitude of BP reductions reported in the literature published in English would reduce the risk of heart disease by  $\sim 22\%$  and stroke by  $\sim 41\%^{62}$  and may lower the BP of adults with hypertension into normal ranges.<sup>63,64</sup> These large and clinically meaningful BP reductions support the use of Tai Ji Quan as an alternative to aerobic exercise as antihypertensive lifestyle therapy based on the literature, whether published in English or Chinese.

Aerobic exercise is the primary exercise modality that professional organizations throughout the world recommend to prevent and treat hypertension.<sup>64</sup> However, less than one-third (28%) of adults with high BP adhere to the current aerobic exercise recommendations for hypertension.<sup>3</sup> Therefore, in order to improve exercise adherence, other exercise options with comparable BP benefits, such as Tai Ji Quan, hold promise as viable alternatives to aerobic exercise, especially when people are unwilling or unable to engage in aerobic exercise to lower their high BP. Indeed, adults with hypertension may show better adherence to Tai Ji Quan than aerobic exercise because it is a safe, low-impact, enjoyable, and inexpensive form of exercise that requires minimal equipment and space.<sup>4-7,65</sup> Tai Ji Quan also provides various other physiological (e.g., managing arthritis symptoms, especially pain) and psychological health benefits (e.g., alleviating fear of falling, reducing stress), some of which (e.g., improving balance) are distinct from more traditional forms of exercise.<sup>8,66</sup>

Our meta-analysis also identified several moderators of the BP response to Tai Ji Quan that warrant further comment. Our most novel finding was that when controlling for other moderators, the magnitude of BP reductions was significantly larger among trials published in Chinese than trials published in English for both SBP ( $\sim$ 17 mmHg vs.  $\sim$ 8 mmHg) and DBP ( $\sim$ 8 mmHg vs.  $\sim$ 3 mmHg). This finding suggests that it is not appropriate to use 1 mean effect size calculation based on trials combined from the English and Chinese literature to represent the efficacy of Tai Ji Quan as antihypertensive lifestyle therapy,<sup>67,68</sup> as all 3 previous meta-analyses did.<sup>14–16</sup> As a result, in our additive regression models, we estimated the greatest potential BP reductions separately for Tai Ji Quan trials published in Chinese and English (Tables 2 and 3). In addition, the larger BP reduction reported among Tai Ji Quan trials published in Chinese than trials published in English emphasizes

that further investigation is needed to explain the large discrepancies in the magnitude of the antihypertensive effects of Tai Ji Quan between trials published in Chinese and English.

Accordingly, we compared differences in the samples, Tai Ji Quan interventions, and study characteristics (Table 4) between Tai Ji Quan trials published in Chinese and English. In relation to sample characteristics, the sole difference we found was that trials published in Chinese had samples with higher DBP at baseline (p = 0.03), which may have contributed to the greater DBP reductions reported.<sup>64</sup> Overall, we did not identify any differences in Tai Ji Quan intervention characteristics that may have contributed to the greater BP reductions reported in trials published in Chinese rather than in English. In relation to the study characteristics, Tai Ji Quan trials published in Chinese differed from trials published in English in that they (1) tended to have smaller sample sizes (p = 0.052), (2) had lower methodological study quality evaluated by using the Downs and Black Checklist (p < 0.001),<sup>28</sup> and (3) had greater publication bias in relation to SBP indicated by the SE of the SBP response to Tai Ji Quan (p=0.048). They also tended to have greater publication bias in relation to DBP, as indicated by the SE of the DBP response to Tai Ji Quan (p = 0.06). These findings indicate that Chinese-language journals are more likely to publish Tai Ji Quan trials showing significant BP reductions, but the trials have noticeable limitations in their study methodology is compared to those published by English-language journals. These limitations likely contributed to the overall greater BP reductions reported in Tai Ji Quan trials published in Chinese.<sup>68,69</sup> We also acknowledge there could be other factors that contributed to the greater BP reductions reported in Tai Ji Quan trials published in Chinese, such as the participants' expectations and the Tai Ji Quan instructors' teaching skills.<sup>70</sup> However, none of these factors were adequately reported to allow for quantitative comparisons.

The 2nd moderator we identified was baseline BP status. When controlling for other moderators, BP reductions were the largest among interventions that involved samples with hypertension (SBP ~14 mmHg/DBP ~6 mmHg), prehypertension (SBP ~11 mmHg/DBP ~5 mmHg), and normal BP (SBP ~8 mmHg/DBP ~3 mmHg).<sup>51</sup> This finding is consistent with the law of initial values,<sup>26</sup> which states that the magnitude of the response is directly related to the initial level of the health outcome being measured.<sup>64</sup>

It should be noted that overall the included Tai Ji Quan trials were of moderate methodological study quality (i.e., 50.1% of items on the Downs and Black Checklist were satisfied<sup>28</sup>). Following the Grading of Recommendations, Assessment, Development, and Evaluation Approach,<sup>71,72</sup> we recommend that readers place moderate certainty in our findings because the included Tai Ji Quan trials exhibited (1) a high/serious risk of bias evaluated by RoB 2.0 and ROBINS-I (Supplementary Tables 5 and 6),

Table 4

Comparisons of the sample, Tai Ji Quan intervention, and study characteristics between Tai Ji Quan trials published in Chinese and English.

	Published in Chinese $(k=13)$		Published in English ( $k = 18$ )	
	k	Mean $\pm$ SD <sup>a</sup>	k	Mean $\pm$ SD <sup>a</sup>
Sample characteristics				
Age (year)	8	$51.9 \pm 14.9$	15	$59.2 \pm 16.3$
Female (%)	10	$78.3 \pm 23.9$	16	$64.0 \pm 18.7$
BMI (kg/m <sup>2</sup> )	6	$24.4 \pm 1.6$	12	$24.6 \pm 1.7$
Baseline SBP (mmHg)	13	$139.1 \pm 16.0$	18	$135.3 \pm 14.8$
Baseline DBP (mmHg)	13	$87.4 \pm 10.3$	18	$80.5 \pm 6.0^{*}$
No Tai Ji Quan practice in the past 6 months (%)	1	7.7	6	33.3
Tai Ji Quan intervention characteristics				
Time (min/session)	12	$47.9 \pm 13.7$	17	$58.4 \pm 4.5^{*}$
Weekly time (min)	12	$210.0 \pm 77.4$	18	$215.3 \pm 90.1$
Total time of intervention (min)	12	$4016.7 \pm 2760.1$	18	$4481.5 \pm 3135.3$
Frequency (sessions/week)	13	$4.6 \pm 1.2$	17	$3.6 \pm 1.5$
Intervention length (week)	13	$21.8\pm25.2$	18	$22.6\pm16.6$
Reported intensity of Tai Ji Quan practice (%)	0	0	1	5.5
Did emphasize movement principles (%)	1	7.7	3	16.7
Did emphasize breathing (%)	4	30.8	2	11.1
Did emphasize relaxation (%)	1	7.7	2	11.1
Study characteristics				
Sample size	13	$73.8 \pm 35.1$	18	$125.8 \pm 87.1$
Publication year	13	$2011.1 \pm 3.8$	18	$2011.4 \pm 5.7$
Methodological study quality (%)	13	$37.0 \pm 10.8$	18	$60.0 \pm 12.6^*$
Adopted RCT design (%)	4	30.8	11	61.1
BP was primary outcome (%)	12	92.3	12	66.7
SE of the SBP response to Tai Ji Quan	13	$0.28\pm0.08$	18	$0.22 \pm 0.08^{*}$
SE of the DBP response to Tai Ji Quan	13	$0.27\pm0.08$	18	$0.22\pm0.08$

Note: <sup>a</sup> For categorical variables, Means and SDs were not applicable; instead, percentages of the number of observations in the identified categories in relation to the total sample (i.e., 13 and 18 for trials published in Chinese and English, respectively) were reported.

\* p < 0.05, based on analysis of variance test for continuous variables and Pearson's  $\chi^2$  test for categorical variables.

Abbreviations: BP = blood pressure; BMI = body mass index; DBP = diastolic blood pressure; RCT = randomized controlled trials; SBP = systolic blood pressure; SE = standard error.

(2) high levels of heterogeneity (i.e.,  $I^2$  ranged from 83.9% to 88.3%, although some heterogeneity was explained with moderators), and (3) some evidence of publication bias, especially in relation to SBP. In addition, we recommend that readers place more certainty in Tai Ji Quan trials published in English than in Chinese for the reasons discussed previously (Table 4).

Although our multiple-moderator models explained a clinically meaningful portion of the variance of the SBP (45.3%) and DBP (65.9%) responses to Tai Ji Quan, it was surprising that these models left a large amount of variance in effects unexplained. We evaluated 20 other potential sources of influence and found that these factors were unrelated to the BP responses to Tai Ji Quan among included trials (Supplementary Table 8, variables tested but not significant as the moderator of the BP response to Tai Ji Quan). Notably, it remains possible that some authors of the included Tai Ji Quan trials inadvertently omitted details of their studies due to space limitations or other unknown reasons, which limited our ability to extract such information and perform analyses. As a result, we acknowledge that additional unidentified moderators may exist in this literature (Supplementary Table 4), such as the intensity of Tai Ji Quan practice (3.2% reported), the emphasis on relaxation (9.7% reported) and breathing techniques (22.7% reported) during Tai Ji Quan practice, the credentials of instructors (48.4% reported various credentials) and whether they corrected the techniques of Tai Ji Quan practice (22.6% reported), the experience of Tai Ji Quan practice among subjects (25.8% reported), adherence to prescribed Tai Ji Quan practice (16.1% reported), medication use at baseline (25.8% reported) and during the intervention (12.9% reported), and maintenance of diet (9.7% reported) and physical activity levels other than Tai Ji Quan practice (9.7% reported) during the intervention. In addition, no intervention (0%) disclosed the time between the last session of Tai Ji Quan and the post-intervention BP measurement. Therefore, it was unclear whether the BP response was confounded by the post-exercise hypotension<sup>21,26</sup> or by detraining.<sup>27</sup> Finally, another limitation of our meta-analysis was that we did not search for the grey literature or Tai Ji Quan trials published in languages other than Chinese and English.

Our meta-analysis also had several strengths. First, we performed meta-regression analysis, allowing us to examine multiple moderators of the BP responses to Tai Ji Quan simultaneously.<sup>18-20</sup> Second, the use of contemporary statistical strategies, in particular the moving constant technique,<sup>44</sup> allowed us to estimate the magnitude of BP reduction at different clinically significant levels of the individual moderators (Tables 2 and 3). Third, although all previous meta-analyses that examined the BP benefits of Tai Ji Quan included trials published in both Chinese and English, ours was the first to examine publishing language as a moderator of the BP response to Tai Ji Quan. Importantly, our results suggest that more confidence should be placed in the Tai Ji Quan trials published in English due to the more noticeable limitations in study methodologies and greater publication bias in the Tai Ji Quan trials published in Chinese.

#### 5. Conclusion

Our new and noteworthy findings are that Tai Ji Quan interventions published in English elicited SBP reductions of 10.4 mmHg and DBP reductions of 4.0 mmHg, while trials published in Chinese elicited SBP reductions of 18.6 mmHg and DBP reductions of 8.8 mmHg among samples with hypertension. Tai Ji Quan may be another viable exercise option to treat hypertension because even the more conservative magnitude of BP reductions among individuals with hypertension reported in the English literature would reduce the risk of cardiovascular disease by up to 40%.<sup>62</sup> The current review also calls attention to the fact that future Tai Ji Quan intervention studies need to (1) improve the reporting of potential moderators of the BP response to Tai Ji Quan and (2) investigate the notable discrepancies we found between trials published in Chinese and English literature.

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#### Authors' contributions

YW participated in the design of the study and contributed to data collection, analysis, and interpretation; BTJ contributed to data analysis and interpretation; SC, YC, and JL contributed to data collection; LSP participated in the design of the study and contributed to data interpretation; all authors contributed to manuscript writing. All authors have read and approved the final version of the manuscript, and agree with the order of presentation of the authors.

#### **Competing interests**

The authors declare that they have no competing interests.

#### Supplementary materials

Supplementary material associated with this article can be found in the online version at doi:10.1016/j.jshs.2020.03.007.

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