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# Resistance training reduces systolic blood pressure in metabolic syndrome: a systematic review and meta-analysis of randomised controlled trials

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## ABSTRACT

**Aim** To evaluate the effects of resistance training on metabolic syndrome risk factors through comparison with a control group.

**Design** Meta-analysis comparing resistance training interventions with control groups. Two independent reviewers selected the studies and assessed their quality and data. The pooled mean differences between resistance training and the control group were calculated using a fixed-effects model.

**Data sources** The MEDLINE, PEDro, EMBASE, SPORTDiscus and The Cochrane Library databases were searched from their earliest records to 10 January 2015.

**Eligibility criteria for selecting studies** Randomised controlled trials that compared the effect of resistance training on metabolic syndrome risk factors with a control group were included. All types of resistance training, irrespective of intensity, frequency or duration, were eligible.

**Results** Only systolic blood pressure was significantly reduced, by 4.08 mm Hg (95% CI 1.33 to 6.82;  $p < 0.01$ ), following resistance training. The pooled effect showed a reduction of 0.04 mmol/L (95% CI  $-0.12$ , 0.21;  $p > 0.05$ ) for fasting plasma glucose, 0.00 (95% CI  $-0.05$ , 0.04;  $p > 0.05$ ) for high-density lipoprotein (HDL) cholesterol, 0.03 (95% CI  $-0.14$ , 0.20;  $p > 0.05$ ) for triglycerides, 1.39 mm Hg (95% CI  $-0.19$ , 2.98;  $p = 0.08$ ) for diastolic blood pressure and 1.09 cm (95% CI  $-0.12$ , 2.30;  $p = 0.08$ ) for waist circumference. Inconsistency ( $I^2$ ) for all meta-analysis was 0%.

**Conclusions** Resistance training may help reduce systolic blood pressure levels, stroke mortality and mortality from heart disease in people with metabolic syndrome.

**Trial registration number** CRD42015016538.

incidence of metabolic syndrome was reduced,<sup>7–10</sup> but in others, resistance training showed no effect on the components of metabolic syndrome.<sup>11–13</sup> Meta-analysis allows the results of independent studies to be grouped.

To the best of our knowledge, the only one previous meta-analysis<sup>14</sup> that has examined the effects of resistance training on metabolic syndrome was published in 2010. The authors included only studies in the English language published up to 2007, with the primary outcomes of HbA<sub>1c</sub> and fat mass percentage. Studies involving four types of intervention were included: resistance training versus control, resistance plus aerobic training versus control, resistance versus aerobic training and resistance training plus diet versus only diet. The systematic review did not investigate the isolated effect of resistance training compared with a control group. The authors did not evaluate the methodological quality of the included studies.<sup>15</sup>

Therefore, the aim of this systematic review and meta-analysis was to evaluate the effects of resistance training on metabolic syndrome risk factors (blood pressure, fasting glucose, triglycerides, high-density lipoprotein (HDL) cholesterol and waist circumference) in comparison with a control group.

## MATERIALS AND METHODS

This systematic review was registered in an international database of systematic reviews in health and social care (registration number CRD42015016538; <http://www.crd.york.ac.uk/PROSPERO/>). The preferred reporting items for systematic review and meta-analyses (PRISMA) guidelines were followed to improve the reporting of this systematic review and meta-analysis.

## Literature search

The MEDLINE, PEDro, EMBASE, SPORTDiscus and The Cochrane Library databases were searched from their earliest records to 10 January 2015, to identify randomised controlled trials that use resistance training as an exercise treatment for metabolic syndrome. The search strategy used a combination of the following Medical Subject Headings (MeSH) terms: randomised controlled trial, metabolic syndrome x and resistance training. Keywords related to these terms were also used (see online supplementary table S1). The reference lists of the included studies were checked to find potential studies that could also be used in this review. There were no language or date of publication restrictions.

## INTRODUCTION

Individuals with metabolic syndrome<sup>1–2</sup> have a greater risk of developing cardiovascular diseases and type 2 diabetes mellitus, and consequently have an increased risk of premature death.<sup>3</sup>

Resistance training is an effective, low-cost strategy to prevent and treat cardiovascular events.<sup>4</sup> There is an inverse association between muscular fitness<sup>5</sup> and strength and the incidence of metabolic syndrome. People who practise resistance training have a 34% lower chance of developing the syndrome.<sup>6</sup>

However, there are conflicting results in papers on the contribution of resistance training and metabolic syndrome disorders. In some trials, the



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## Study selection

Study selection was performed by two independent researchers (IRL and SNL), with disagreements resolved by consensus. If necessary, a third researcher (JNJ) was consulted. Only randomised controlled trials that compared resistance training with a control group (no intervention) were included in this review. Studies that used diet intervention were included if this intervention was equal for all the groups in the study. Trials were eligible if they included participants with metabolic syndrome and assessed the components of the syndrome: elevated fasting plasma glucose, triglycerides, HDL-cholesterol, blood pressure and waist circumference. As the aim of the study was to evaluate the effect on all metabolic syndrome risk factors, studies that did not evaluate at least four of these outcomes were excluded. All types of resistance training, irrespective of intensity, frequency or duration, were eligible for inclusion.

## Data extraction and risk of bias assessment

For risk of bias assessment, two different tools were used, the PEDro scale<sup>16 17</sup> and the Grading of Recommendations Assessment, Development and Evaluation (GRADE).<sup>18 19</sup> The PEDro scale is a valid measure of the methodological quality of clinical trials and is based on the Delphi list developed by Verhagen *et al.*<sup>20</sup> The GRADE approach evaluates the quality of evidence for outcomes (meta-analysis) reported in systematic reviews. In addition, it is a systematic and explicit approach that allows judgements to be made about strength of evidence and is an effective method for linking evidence quality and clinical recommendations.

Two reviewers (IRL and SNL) independently assessed the risk of bias of individual studies using the PEDro scale.<sup>16 17</sup> If trials were already listed on the PEDro database (<http://www.pedro.org.au/>), these scores were adopted. A PEDro score of 7 or greater was considered of 'high quality', studies with a score of 5 or 6 were considered of 'moderate quality' and those with a score of 4 or less were deemed of 'poor quality'.<sup>21–23</sup> Any disagreements in the scoring of trials were resolved consensually. Methodological quality was not an inclusion criterion. Two reviewers (IRL and SNL) also independently extracted outcome data using a standardised data extraction form.

The GRADE approach was used by two independent reviewers (IRL and SNL) to evaluate the overall quality of evidence and the strength of the recommendation,<sup>18 19</sup> as advocated by the Cochrane Back Review Group.<sup>24</sup> The overall quality of evidence was initially regarded as 'high' but was downgraded by one level for each of the three factors encountered: risk of bias (>25% of participants from studies with low-quality methods—PEDro score <7 points), inconsistency of results (substantial  $I^2$  statistic) and imprecision (<400 participants in total for each outcome). Publication bias assessment with a funnel plot was not performed and indirectness was not considered for this review owing to the presence of a specific population, relevant outcome measures and direct comparisons.

The following factors were used to define the quality of evidence: high quality—further research is unlikely to change our confidence in the estimate of effect; moderate quality—further research is likely to have an important impact on our confidence in the estimate of effect and might change the estimate; low quality—further research is likely to have an important impact on our confidence in the estimate of effect and is likely to change the estimate; and very low quality—we are uncertain about the estimate.

Extracted data included final values of means, SDs and sample sizes of triglycerides, fasting glucose, HDL-cholesterol,

blood pressure and waist circumference. When final values were not available, change scores were used. When there was insufficient information, the authors were contacted. As SD values are not always reported by researchers, where necessary, these data were imputed or calculated using methods recommended in the Cochrane Handbook for Systematic Reviews of Interventions. As an example, in the study by Castaneda,<sup>25</sup> SD was obtained from the SE of the mean multiplied by the square root of the sample size.

Blood pressure values were expressed in millimetre of mercury, waist circumference in centimetres and glucose, triglycerides and HDL-cholesterol in millimole per litre. Where appropriate, data were converted to these units of measurement.

## Statistical analysis

Pooling was carried out using Comprehensive Meta-Analysis software, V2.2.064 (BioStat, Englewood, New Jersey, USA). When trials were sufficiently homogeneous (ie, an  $I^2$  value <50%), pooled effects were calculated using a fixed-effects model, whereas random effects were used to estimate the pooled effects of heterogeneous trials (ie, an  $I^2$  of 50% or more). Mean difference (MD) with 95% CIs was calculated.

In addition, a stratified exploratory analysis was performed using the same procedures as the main analysis, comparing high (PEDro  $\geq 7$ ) versus low/moderate (PEDro  $\leq 6$ ) methodological quality, and short-term (<6 months) versus long-term ( $\geq 6$  months) interventions, as used in a previous study.<sup>26</sup>

## RESULTS

### Description of included studies

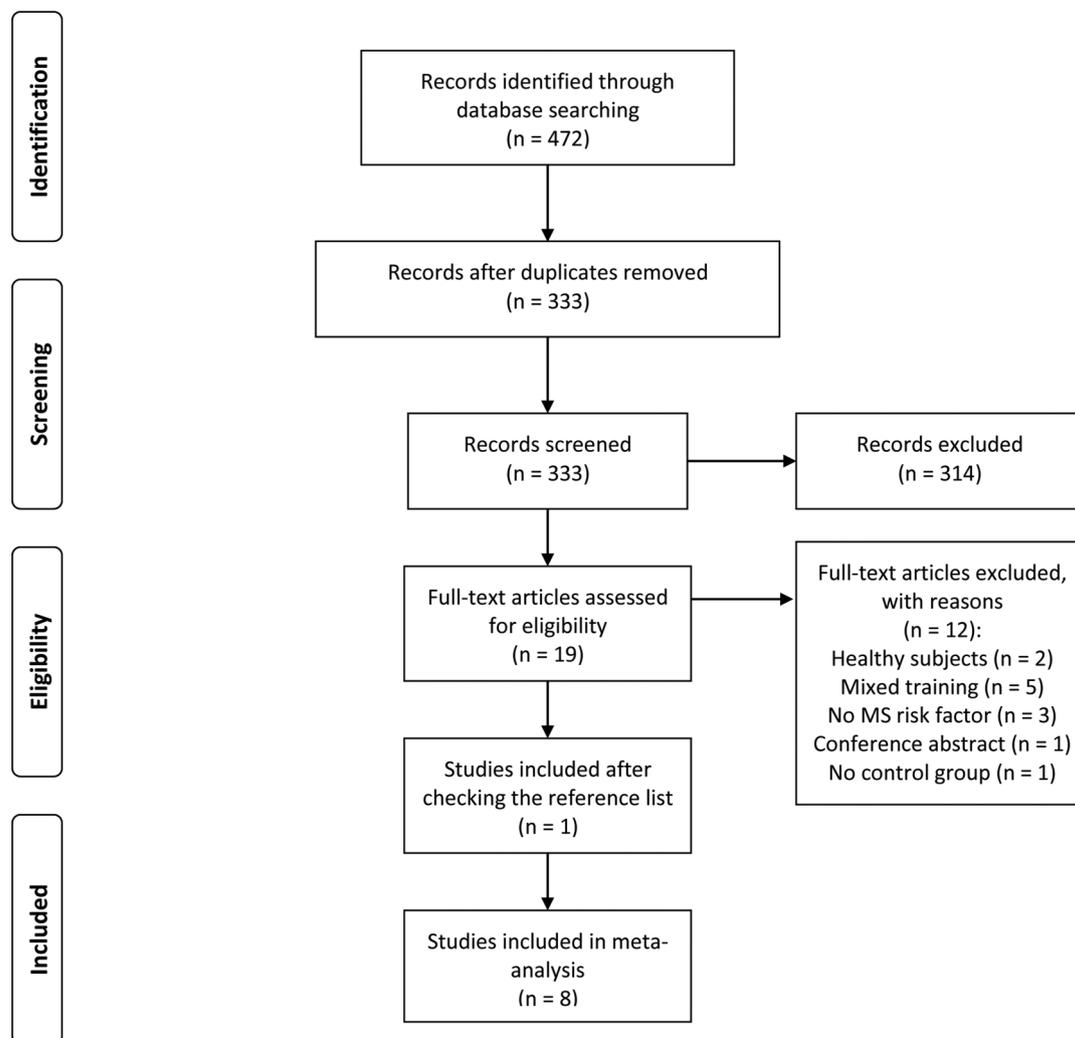
After the removal of duplicates, the search strategy identified 333 titles. Screening of titles and abstracts identified 19 potentially eligible articles, and 7 original trials were included.<sup>7 8 10 11 25 27 28</sup> A further study<sup>29</sup> was included after checking the reference lists of included trials. The reasons for excluding articles were: use of healthy subjects,<sup>30 31</sup> intervention with mixed training, that is, resistance plus aerobic training,<sup>32–36</sup> no evaluation of metabolic syndrome risk factors,<sup>37–39</sup> conference abstract<sup>40</sup> and no control group<sup>13</sup> (figure 1).

A total of 341 men and 178 women were included in the meta-analysis. Three studies included only men,<sup>10 11 28</sup> and five included a mixed sample of men and women, with 64.5% (n=40),<sup>25</sup> 44.8% (n=13),<sup>27</sup> 36.2% (n=46),<sup>29</sup> 39.5% (n=8)<sup>8</sup> and 62.2% (n=71)<sup>7</sup> of women, respectively. The training period ranged from 12 weeks to 9 months, and all studies had an incremental workload in intensity or volume. Online supplementary table 2 shows the characteristics of the included trials.

### Resistance training as a treatment of metabolic syndrome

A fixed-effects model was used for all meta-analyses. The results of meta-analysis comparing the effects of resistance training with a control group show that resistance training is significantly superior to control groups in terms of reducing systolic blood pressure (seven studies, n=476,  $I^2=0\%$ , MD 4.08 mm Hg (95% CI 1.33 to 6.82)).

However, the results of pooling data show that resistance training is not superior to control interventions (no intervention) in improving fasting plasma glucose (seven studies, n=370,  $I^2=0\%$ , MD 0.04 mmol/L (95% CI -0.12 to 0.21)), HDL-cholesterol (eight studies, n=497,  $I^2=0\%$ , MD 0.00 mmol/L (95% CI -0.05 to 0.04)), triglycerides (eight studies, n=497,  $I^2=0\%$ , MD 0.03 mmol/L (95% CI -0.14 to 0.20)), diastolic blood pressure (seven studies, n=476,  $I^2=0\%$ , MD 1.39 mm Hg (95% CI -0.19, 2.98)) and waist



**Figure 1** Flow chart of studies included.

circumference (eight studies,  $n=497$ ,  $I^2=0\%$ , MD 1.09 cm (95% CI  $-0.12$  to 2.30)) (figures 2 and 3).

### Methodological quality

One study<sup>29</sup> was considered ‘high quality’, three studies<sup>8 25 28</sup> were considered ‘moderate quality’ and four studies<sup>7 10 11 27</sup> were considered ‘poor quality’. All included trials had random allocation between group comparisons and provided points and estimates of variability. Concealed allocation was performed in two studies.<sup>8 29</sup> Owing to the nature of the interventions, blinding of participants and therapists was not possible. Assessor blinding was implemented in two of included studies.<sup>25 29</sup> In addition, four studies had adequate follow-up,<sup>8 25 28 29</sup> and two studies included an intention-to-treat analysis.<sup>25 29</sup> Complete details are reported in online supplementary table 3.

### Secondary exploratory analysis

Exploratory analysis was performed comparing short-term (<6 months) versus long-term ( $\geq 6$  months) studies (see online supplementary appendix figure 1–appendix figure 6), and low/moderate methodological quality (PEDro  $\leq 6$ ) versus high methodological quality (PEDro  $\geq 7$ ) studies (see online supplementary appendix figure 7–appendix figure 11). Stratified analysis by methodological quality for fasting glucose results was not possible as the high-quality study<sup>29</sup> did not report this outcome.

Long-term trials are more effective than short-term trials in reducing systolic blood pressure (four studies,  $I^2=0\%$ , MD 3.85 mm Hg (95% CI 0.55 to 7.14);  $p=0.02$ ). There was also a reduction in diastolic blood pressure, although this was not statistically significant (four studies,  $I^2=0\%$ , MD 1.97 mm Hg (95% CI  $-0.20$  to 4.14);  $p=0.07$ ). Other potential influences of these aspects were not detected as comparisons of subgroups revealed no differences in pooled estimates with overlapping CIs.

No inference could be made about the comparison between low/moderate-quality and high-quality studies because of the small number of high-quality studies.

### GRADE assessment of main analyses

On the basis of the GRADE system (see online supplementary table 4), pooled data of HDL-cholesterol, triglycerides, systolic blood pressure, diastolic blood pressure and waist circumference were classified as moderate-quality evidence. All these variables were downgraded one level owing to the presence of risk of bias (more than 25% of participants from studies of low/moderate methodological quality, PEDro score  $< 7$  points). The evidence for fasting plasma glucose lost points owing to the risk of bias and imprecision (fewer than 400 participants in the meta-analysis) and was classified as low-quality evidence.

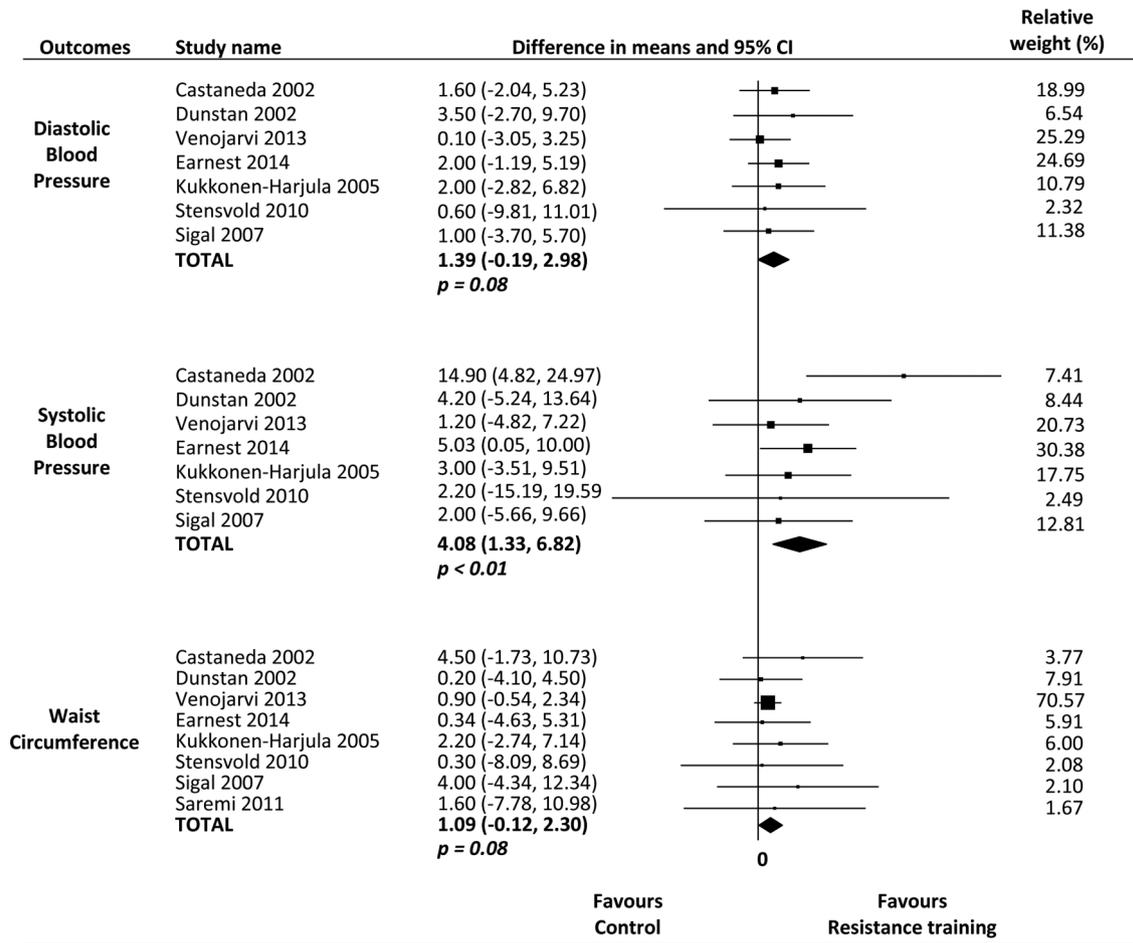


Figure 2 Effects of resistance training on clinical parameters of metabolic syndrome.

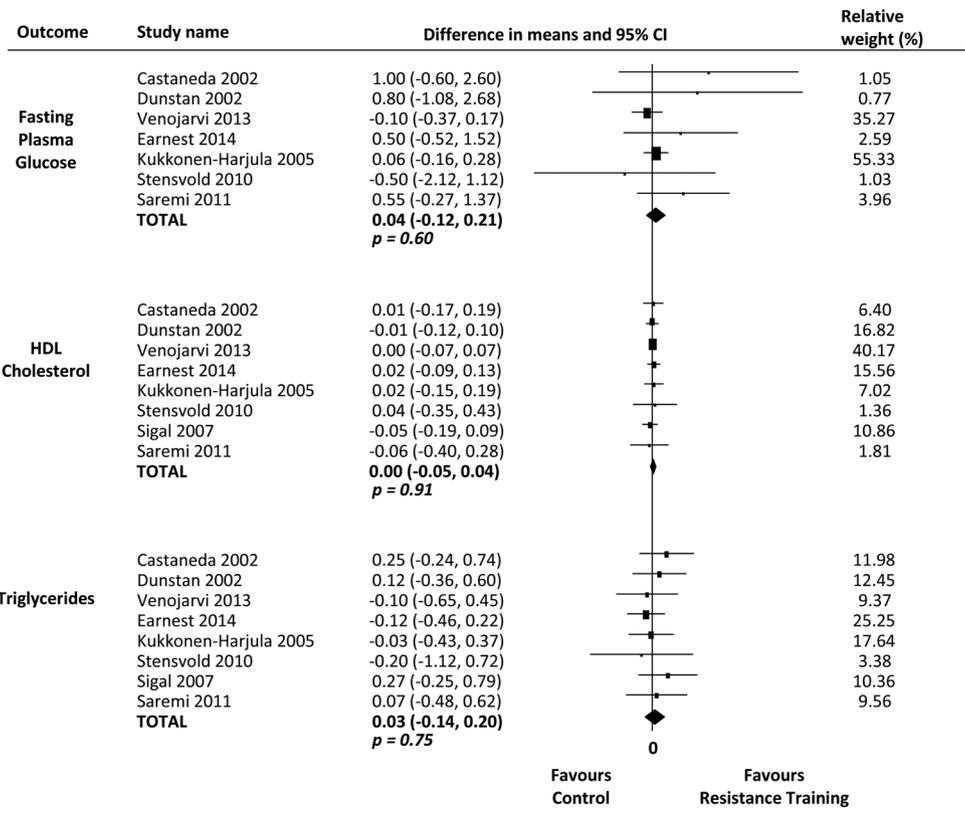


Figure 3 Effects of resistance training on metabolic parameters of metabolic syndrome.

## DISCUSSION

This systematic review provides evidence of moderate quality that resistance training reduces systolic blood pressure in adults with metabolic syndrome. Although there seems to be a trend that resistance training slightly improves diastolic blood pressure and waist circumference, the meta-analysis may have been underpowered for detection of differences in these two risk factors. Resistance training had no effect on metabolic parameters, that is, fasting glucose, HDL-cholesterol and triglycerides.

### Systolic blood pressure reduced by resistance training

There was a reduction in systolic blood pressure of 4.1 mm Hg, which extends findings in previous studies that reported reductions after resistance training intervention.<sup>14 41 42</sup> Lewington *et al*<sup>43</sup> found that even a reduction of up to 2 mm Hg can reduce stroke mortality by around 10% and mortality from heart disease by about 7%. Although these reductions seem proportionately small, it is related to changes in only one risk factor, systolic blood pressure.

Thus, the present study reinforces the importance of physical training as an important strategy in the treatment and prevention of this syndrome and shows that further randomised controlled trials are required to conclusively determine the effect of resistance training on the parameters of metabolic syndrome. Moreover, the methodological quality of future studies should be improved. Researchers should be concerned mainly to concealed allocation, blinding of assessors and the inclusion of an intention-to-treat analysis.

### Comparison with other studies

Regarding metabolic outcomes, the results extend those of Cornelissen *et al*.<sup>42</sup> The authors did not find any significant changes in fasting glucose, HDL-cholesterol and triglyceride values. When the studies were considered individually, only one found significant positive changes in HDL-C<sup>28</sup> and one showed improvement in triglyceride and fasting plasma glucose values.<sup>10</sup> Both studies used only male subjects. Whereas five of the studies included in this review comprised volunteers of both sexes, and metabolic changes caused by menopause may influence the behaviour of these risk factors,<sup>44</sup> further research is needed with specific populations to better understand these effects.

One interesting point to consider in this study is the consistency found in the meta-analyses. Such consistency is probably due to the similarity of the effect sizes. A specific population of similar age, affected by the same condition (metabolic syndrome), and undergoing similar interventions may have contributed to this lack of heterogeneity. However, it is important to note that the absence of heterogeneity, as measured by  $I^2$ , does not imply homogeneity.

Considering these results, we propose periodised and individualised resistance training as part of treatment to reduce systolic blood pressure in patients with metabolic syndrome.

### Strengths and limitations

The strengths of this systematic review are its search protocol and the inclusion of only randomised controlled trials. In addition, trials published in any language were eligible for inclusion. Another strength of this review is that the training effect was explored through quantitatively pooled trials. Inconsistency ( $I^2$ ) was assessed to evaluate the consistency of the results. It is important to show that the variation in findings is compatible with chance alone. The  $I^2$  of the meta-analysis was 0% for all

risk factors. Also, the overall quality of the evidence was assessed using the GRADE approach, which evaluates quality of evidence and is an effective method for linking evidence quality and clinical recommendations.

A limitation of this review is that publication bias was not assessed with a funnel plot, as tests for funnel plot asymmetry should be used only when there are at least 10 studies included in the meta-analysis.<sup>24</sup> Another limitation, not of this study particularly, is the low number of participants composing this meta-analysis. It shows that, despite the advance of exercise science in recent years, there is a lack of randomised trials evaluating these conditions (metabolic syndrome and resistance training).

In summary, resistance training is an effective treatment for reducing systolic blood pressure levels by approximately 4 mm Hg in people with metabolic syndrome. This effect size is clinically meaningful because it would translate to a 10% reduction in stroke mortality and 7% reduction in deaths from heart disease if achieved at a population level.

### What is already known?

- ▶ Resistance training is an effective and low-cost method to prevent and treat cardiovascular diseases.
- ▶ There is an inverse association between muscular strength and the incidence of metabolic syndrome.

### What are the new findings?

- ▶ Resistance training reduces systolic blood pressure by approximately 4.1 mm Hg.
- ▶ At the society level, a reduction of up to 2 mm Hg can reduce stroke mortality by around 10% and mortality from heart disease by about 7%.
- ▶ There was no statistical difference between resistance training and control in reducing diastolic blood pressure and waist circumference.

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