

# NIH Public Access

Author Manuscript

J Womens Health (Larchmt). Author manuscript; available in PMC 2008 July 9

J Womens Health (Larchmt). 2004 December; 13(10): 1148–1164.

## Aerobic Exercise and Lipids and Lipoproteins in Women: A Meta-Analysis of Randomized Controlled Trials

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## Abstract

**Background**—Cardiovascular disease (CVD) in women is the leading cause of mortality in the United States, and less than optimal lipid and lipoprotein levels are major risk factors for CVD. The purpose of this study was to use the meta-analytic approach to examine the effects of aerobic exercise on lipids and lipoproteins in women.

**Methods**—Studies were retrieved via computerized literature searches, review of reference lists, hand searching selected journals, and expert review of our reference list. The inclusion of studies was limited to randomized controlled trials published in the English language literature between January 1955 and January 2003 in which aerobic exercise was used as the primary intervention in adult women aged  $\geq 18$  years. One or more of the following lipids and lipoproteins were assessed: total cholesterol (TC), high-density lipoprotein cholesterol (HDL-C), low-density lipoprotein cholesterol (LDL-C), and triglycerides (TG).

**Results**—Using a random effects model, statistically significant improvements were observed for all lipids and lipoproteins (TC,  $\vec{X} \pm$  SEM, -4.3 ± 1.3 mg/dl, 95% CI -6.9 to -1.7 mg/dl; HDL-C,  $\vec{X} \pm$  SEM, 1.8 ± 0.9 mg/dl, 95% CI 0.1 to 3.5 mg/dl; LDL-C,  $\vec{X} \pm$  SEM, -4.4 ± 1.1 mg/dl, 95% CI -6.5 to -2.2 mg/dl; TG,  $\vec{X} \pm$  SEM, -4.2 ± 2.1 mg/dl, 95% CI -8.4 to -0.1 mg/dl). Reductions of approximately 2%, 3%, and 5%, respectively, were observed for TC, LDL-C, and TG, whereas an increase of 3% was observed for HDL-C.

**Conclusions**—Aerobic exercise is efficacious for increasing HDL-C and decreasing TC, LDL-C, and TG in women.

## Introduction

In women, cardiovascular disease (CVD) is the leading cause of mortality in the United States. Recent data indicate that more than 360,000 women die annually from diseases of the heart. <sup>1</sup> Less than optimal lipid and lipoprotein levels, especially low-density lipoprotein cholesterol (LDL-C), are major risk factors for CVD.<sup>2</sup> One lifestyle approach that may improve lipid and lipoprotein concentrations in women is aerobic exercise, a cost-effective, nonpharmacological intervention that is available to the vast majority of the general public.<sup>2</sup> However, previously published randomized controlled trials investigating the effects of aerobic exercise on lipids and lipoproteins in women have shown less than overwhelming results.<sup>3–43</sup> For example, of the previously cited studies, statistically significant decreases were reported for only 19% of total cholesterol (TC) outcomes, 11% for LDL-C outcomes, 18% for high-density lipoprotein

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cholesterol (HDL-C) outcomes, and 8% for triglyceride (TG) results.<sup>3–43</sup> In addition, although a meta-analysis that focused specifically on the effects of exercise on lipids and lipoproteins in women has been published,<sup>44</sup> this study was conducted more than 15 years ago and was limited to studies published through December 1987. A total of 27 randomized and nonrandomized controlled trials as well as studies without a control group were included in that analysis. Across all designs and categories, a statistically significant reduction of 4 mg/dl (2%) was observed for TC, and a reduction of 9 mg/dl (10%) was observed for TG. No statistically significant differences were observed for LDL-C or HDL-C. Since that time, many additional studies on this topic have been published or located.<sup>3,4,6–15,17–19,21–31,33–38</sup>, <sup>40,41,43</sup> The Cochrane Collaboration, an international organization dedicated to the conduct of systematic reviews in healthcare, suggests that because results on a given topic can change over time, systematic reviews usually should be updated on a yearly basis in order to have the most recent information available on a chosen topic.<sup>45</sup> In addition, more sophisticated techniques for handling meta-analysis datasets have been developed (e.g., use of the random effects model) as well as a greater reliance on limiting the inclusion of studies to randomized controlled trials.<sup>45</sup> Furthermore, we have reported the lack of statistically significant findings from the individual studies cited  $^{3-43}$  using a vote-counting approach (percentage of significant findings) that is less accurate than using the actual values reported in the studies.<sup>45</sup> Metaanalysis is an approach in which statistical techniques are used to integrate the results of studies meeting predetermined inclusion criteria.<sup>45</sup> The strengths of meta-analysis include the ability to (1) increase the power of primary endpoints, (2) increase estimates of treatment effectiveness, and (3) resolve discrepancies when studies disagree.

Thus, given (1) the negative effect of less than optimal lipid and lipoprotein levels on CVD in women, (2) the potential for aerobic exercise to have a positive effect on lipids and lipoproteins in women, (3) the less than overwhelming results of previous studies dealing with the effects of aerobic exercise on lipids and lipoproteins in women, (4) the absence of any recent meta-analytic work dealing with the effects of aerobic exercise on lipids and lipoproteins in women, and (5) the strengths of meta-analysis, the purpose of this study was to use the meta-analytic approach to examine the effects of aerobic exercise on lipids and lipoproteins in women.

#### **Materials and Methods**

#### Data sources

Studies for this meta-analysis were retrieved via (1) electronic databases (MEDLINE, EMBASE, Current Contents, Dissertation Abstracts International), (2) review of the reference lists from review articles and original studies, (3) hand searching selected journals, and (4) review of our reference list by an expert on lipids and lipoproteins (Dr. William Haskell, personal communication, May 7, 2003). Keywords used in our electronic database searches included various combinations of terms including, but not limited to, the following: exercise, physical fitness, physical activity, aerobic, cholesterol, lipids, lipoproteins, adults, humans. Hand searching of selected journals was performed because not all trial reports are included in electronic databases, and even when they are included, they may not be indexed with terms that allow them to be identified as trials that meet the meta-analysis inclusion criteria.<sup>45</sup>

#### **Study selection**

The inclusion criteria for this study were as follows: (1) randomized controlled trials, (2) chronic aerobic exercise of at least 8 weeks as an intervention, (3) adult females aged  $\geq$ 18 years, (4) studies published in journal, dissertation, or master's thesis format, (5) studies published in the English language, (6) studies published between January 1955 and January 2003, and (7) one or more of the following lipids and lipoproteins assessed in the fasting state: TC, HDL-C, LDL-C, and TG. Studies published in foreign languages were not included

because it was beyond the scope of this project. We used 1955 as a starting point for our search because this appears to be the first year in which a controlled trial addressing the effects of exercise on lipids and lipoproteins was conducted.<sup>46</sup> We included only those groups that met our inclusion criteria from each study. For example, if a study included two intervention groups (aerobic exercise only, aerobic exercise plus diet) and a control group, we included the aerobic exercise and control groups but not the aerobic exercise plus diet group.

#### **Data abstraction**

A coding form was developed that could hold more than 400 items from each study. The major categories of items that were coded included (1) study characteristics (e.g., year and source), (2) physical characteristics of subjects (e.g., age and height), (3) lipid assessment characteristics (e.g., position and number of hours fasted), (4) primary outcomes (TC, HDL-C, LDL-C, and TG), and (5) secondary outcomes (body weight, body mass index [BMI] in kg/m<sup>2</sup>, percent body fat, and maximum oxygen consumption ( $V_{o_2}max$ ) in ml/kg<sup>-1</sup>/min<sup>-1</sup>). All studies were coded by the first two authors, independent of each other. We then reviewed every data point for accuracy and consistency. Interrater agreement for the initial review of studies using Cohen's kappa statistic<sup>47</sup> was 0.95. Any discrepancies were resolved by consensus. If no consensus could be reached, the third author served as an arbitrator.

#### Statistical analysis

Primary and secondary outcomes—The primary outcomes in this study were changes in TC, HDL-C, LDL-C, and TG, and secondary outcomes included body weight, BMI, percent body fat, and  $V_{02}$  max. As all studies used a parallel-group design, changes in primary and secondary outcomes were calculated as the difference (exercise minus control) of the changes (initial minus final) in these mean values. Variances for each group (exercise and control) were calculated from variances at baseline and final measurement using standard methods. 48 Pooled outcomes for primary and secondary outcomes were calculated by assigning weights equal to the inverse of the total variance for net changes in these outcomes. A random effects model was used for all analyses, <sup>49,50</sup> and 95% confidence intervals (95% CI) were used to establish the precision of our results. If the CI did not cross zero, our results were considered to be statistically significant. Statistical heterogeneity of study results was examined using the Q statistic.<sup>51</sup> An alpha level of  $p \le 0.10$  vs.  $p \le 0.05$  was used to determine statistically significant heterogeneity because this test tends to suffer from low power.<sup>52</sup> However, as the Q statistic can give misleading results because of issues surrounding power, we also used a recently developed statistic (I<sup>2</sup>) that measures the degree of inconsistency across studies in a metaanalysis.<sup>52</sup> This statistic, which is based on the Q statistic, ranges from 0% to 100%, with larger numbers representing greater inconsistency. Values of 25%, 50%, and 75%, respectively, are indicative of low, moderate, and high degrees of inconsistency.<sup>52</sup> Publication bias was examined using the procedure of Egger et al.<sup>53</sup> Study quality was examined using a previously validated (face validity) and reliable (r = 0.77) three-item questionnaire designed to assess bias, specifically randomization, blinding, and withdrawals/dropouts. The minimum number of points possible ranges from a low of 0 to a high of 5, with higher numbers representing greater study quality.<sup>54,55</sup> In addition to our overall results for TC, LDL-C, HDL-C, and TG, we also ran our analyses with those groups in which one or more subjects were taking hormone replacement therapy (HRT) deleted from the model, groups in which changes in diet occurred deleted from the model, and the one study in which all subjects were hyperlipidemic deleted from the model.

**Subgroup analyses**—Clinical heterogeneity for TC, HDL-C, LDL-C, and TG was examined by performing *a priori* subgroup analyses for categorical variables using random effects ANOVA models (method of moments approach), where Q<sub>b</sub>, which is similar to the Fratio, represents between-group differences.<sup>56</sup> Variables that were examined included study

source, country, menopausal status (pre vs. post), use of drugs that might affect lipids and lipoproteins (yes vs. no), race/ethnicity, cigarette smoking, diabetes, overweight/obesity, position of lipid assessment, and training modality. We did not conduct subgroup analyses for alcohol consumption because none of the studies reported the inclusion of subjects in which all or none of the subjects consumed alcohol. In addition, we did not conduct subgroup analyses for diet or physical activity because only three studies reported that some subjects were physically active prior to taking part in the study.<sup>3,8,23</sup> or that some of the subjects had changes in diet during the study that could affect lipid and lipoprotein levels.<sup>23,40,43</sup> Subgroup analysis according to hyperlipidemic status or the presence of CVD was also not possible because only one study reported that all subjects were hyperlipidemic,<sup>30</sup> and none reported that all women had CVD. Because of the multiple subgroup analyses we conducted, our alpha value for statistical significance was set at  $p \leq 0.01$ .

**Regression analysis**—Clinical heterogeneity for TC, HDL-C, LDL-C, and TG was examined by performing *a priori* simple, weighted, generalized least-squares meta-regression (random effects, method of moments approach) for continuous variables.<sup>56</sup> Variables that were regressed included initial lipid level, age, height, initial as well as treatment effect changes in body weight, BMI, percent body fat, and  $V_{o_2}max$ , number of hours fasted prior to the assessment of lipids and lipoproteins, number of hours that exercise was avoided prior to the assessment of lipids and lipoproteins, length, frequency, intensity, duration, and total minutes of training (length × frequency × duration), and compliance to the exercise protocol. We did not perform any type of multiple regression because of missing data for different variables from different studies and our desire to include as many data as possible for each analysis. Because of the large number of simple regression analyses we conducted, we used 99% vs. 95% CIs to establish statistical significance. If the CI did not cross zero, we considered our results to be statistically significant.

**Data reporting**—Descriptive statistics are reported as mean  $\pm$  standard deviation ( $X \pm$  SD), and results of analysis (primary and secondary outcomes) are reported as mean  $\pm$  standard error of the mean ( $\overline{X} \pm$  SEM). Relative (percent) changes in primary and secondary outcomes were calculated by subtracting the change outcome (final minus initial) from the initial value for each group (exercise vs. control) and then multiplying by 100. The relative (percent) change reported was calculated by subtracting the percent change in the exercise group from the percent change in the control group.

#### Results

#### Study characteristics

**Studies included**—Of the 3750 citations reviewed, 41 studies met our inclusion criteria<sup>3–43</sup> (Table 1).

**Country of origin**—Twenty-nine of the 41 studies (71%) were conducted in the United States, 5,7,8,10–12,14–18,20–22,24–29,32–36,39–41,43 and two each were conducted in Canada, <sup>30,31</sup> Japan, <sup>13,37</sup> and the United Kingdom.<sup>4,9</sup> One study each was conducted in Australia, <sup>23</sup> Finland, <sup>3</sup> Kuwait, <sup>6</sup> Sweden, <sup>42</sup> Switzerland, <sup>38</sup> and Turkey.<sup>19</sup>

**Study design**—All included studies used a parallel-group design. Only two studies appeared to use the intention-to-treat approach in the analysis of their data.<sup>13,38</sup> A total of 98 groups (56 exercise, 42 control) representing 1715 subjects (1022 exercise, 693 control) and up to 56 outcomes were available for pooling. The number of exercise groups exceeded the number of control groups because some studies had more than one exercise group. The percentage of subjects who were not available for follow-up assessment ranges from 0% to 66.7% in the

exercise groups ( $\overline{X} \pm$  SD, 22.6 ± 17.3%) and 0% to 40.0% in the control groups ( $\overline{X} \pm$  SD, 13.8 ± 13.9%). Study quality ranged from 0 to 5 (median = 1).

#### Subject characteristics

**Race**—Baseline characteristics of the subjects are shown in Table 2. For those subjects who reported information on race/ethnicity, three studies reported that all subjects were white,<sup>7,8, 29</sup> two reported that all subjects were Japanese,<sup>13,37</sup> and one reported that all subjects were Hispanic.<sup>21</sup> One study each reported that subjects consisted of whites, blacks, and Hispanics, <sup>10</sup> primarily whites (89%),<sup>22</sup> whites, African Americans, and Hispanics,<sup>33</sup> or Caucasians (89%) and one Asian.<sup>34</sup>

**Menopausal status/HRT**—Nine studies reported that all subjects were premenopausal,<sup>3</sup>, 10,17,18,21,24,27,33,38 and another ten reported that all subjects were postmenopausal.<sup>8</sup>, 12,22,25,29–31,34,36,40 Another four studies included both premenopausal and postmenopausal women and reported data for both groups combined.<sup>4,13,23,37</sup> One other study included both premenopausal and postmenopausal women but reported data separately for each group.<sup>6</sup> For those studies that included data on groups of postmenopausal women, seven reported that none of the subjects were taking any type HRT,<sup>8,22,25,30,31,34,40</sup> and the other three reported one or more subjects were taking some type of HRT.<sup>12,29,36</sup> We were unable to calculate the percentage of women taking HRT because of a lack of reported data.

**Medication**—Fourteen studies reported that none of the subjects were taking any type of medication that could affect lipids and lipoproteins, 4,6,8,18,20,22,24,25,29-31,34,37,40 whereas ten studies reported that some of the subjects were taking some type of drug that could affect lipids and lipoproteins. 10-12,17,21,23,32,33,36,38 Of these ten studies, two (20%) were limited to postmenopausal women. 12,36

**Cigarette smoking/alcohol**—Eighteen studies reported that none of the subjects smoked cigarettes, 4,10–12,14,15,17,18,24,26,28–31,33,34,38,40 and six reported that some of the subjects smoked. 8,13,22,23,41,42 In relation to alcohol consumption, eight studies reported that some of the subjects consumed alcohol. 5,8,13,15,17,23,31,38

**Diet/physical activity**—Eighteen studies reported that there were no changes in diet that could affect lipid and lipoprotein levels, 5,8,10-13,18,22,24,26,29-34,36,41 and two studies reported that some of the subjects had changes in diet that could affect lipid and lipoprotein levels.<sup>23,40</sup> One other study reported that there were no changes in diet that could affect lipid and lipoprotein levels in the exercise group but that changes in diet that could affect lipids and lipoproteins did occur in the control group.<sup>43</sup> Thirty-six studies reported that none of the subjects were physically active prior to taking part in the study, 4-7,9-12,14-22,24-35,37-43 two reported that some of the subjects were physically active, 8,23 and one reported that all

<sup>45</sup> two reported that some of the subjects were physically active,<sup>8,25</sup> and one reported that all the subjects were physically active prior to taking part in the study.<sup>3</sup> One other study reported that none of the subjects in the exercise group but some of the subjects in the control group were physically active prior to taking part in the study.<sup>13</sup>

**Comorbidities**—Seven studies reported that all the subjects were overweight/obese,  $^{14-18}$ ,  $^{21,28,40}$  and another two studies reported that all subjects had diabetes, one with non-insulindependent diabetes mellitus<sup>40</sup> and another with insulin-dependent diabetes mellitus.  $^{42}$  One study reported that all subjects were hyperlipidemic,  $^{30}$  and none reported that all subjects had CVD.

#### Lipid assessment characteristics

Five studies reported assessment of lipids and lipoproteins in the sitting position, 6,15,17,31,  $3^{2}$  and another five studies reported assessment in the supine position. 3,5,11,16,33 A total of eight studies reported the assessment of lipids and lipoproteins at least twice during each testing period. 10,11,24,30-33,36 The number of hours that subjects fasted prior to the assessment of lipid and lipoproteins ranged from 9 to 14 hours ( $X \pm$  SD,  $11.9 \pm 0.95$  hours), and the number of hours that exercise was avoided prior to the assessment of lipids and lipoproteins ranged from 12 to 72 hours ( $X \pm$  SD,  $30.4 \pm 19.1$  hours). Twenty-two studies reported that they calculated LDL-C, 3,5,6,10,12,16,17,18,19,21,24-26,28-33,36,42,43 and one reported the direct estimation of LDL-C. 13

#### **Training program characteristics**

A description of the training program characteristics is shown in Table 3. Twenty-four groups from 19 studies (approximately 43%) used walking as the primary training modality, <sup>4,8–10</sup>, <sup>12,13,17,18,21,27–35,40</sup> six groups from four studies used aerobic dance, <sup>6,14,19,41</sup> five groups from three studies used jogging, <sup>5,32,39</sup> two groups from two studies used cycling, <sup>42</sup>, <sup>43</sup> and one group from one study used stair climbing. <sup>43</sup> Another six groups from six studies participated primarily in a combination of walking and jogging, <sup>7,16,20,36,38,43</sup> five groups from three studies participated in walking, jogging, and cycling, <sup>15,22,26</sup> three groups from three studies participated in miscellaneous types of activities, <sup>11,37,43</sup> and one group from one study each participated primarily in walking and aerobic dance<sup>23</sup> or walking and cycling. <sup>25</sup> As can be seen, compliance data were available for only 55% of the groups.

#### Primary outcomes

Results for primary outcomes are shown in Table 4 and Figure 1.

**TC**—Across all categories, a statistically significant reduction of approximately 2% was observed. This reduction represents the difference between the pre to post changes in the exercise and control outcomes. No statistically significant publication bias was observed (p =0.37). This suggests that our results were not biased toward positive results. With those groups in which one or more subjects were taking HRT deleted from the model, decreases in TC continued to be statistically significant ( $-4.4 \pm 1.4 \text{ mg/dl}$ , 95% CI -7.2 to -1.6 mg/dl). In addition, changes in TC remained statistically significant when we deleted the study in which all subjects were hyperlipidemic  $(-4.1 \pm 1.4 \text{ mg/dl}, 95\% \text{ CI} - 6.8 \text{ to} - 1.4 \text{ mg/dl})$  as well as those groups in which changes in diet occurred ( $-4.4 \pm 1.4 \text{ mg/dl}, 95\% \text{ CI} - 7.1 \text{ to} - 1.7 \text{ mg/}$ dl). Furthermore, no statistically significant or clinically relevant differences were observed for those variables in which subgroup analyses were performed, including differences in TC between premenopausal and postmenopausal women ( $Q_b = 0.03$ , p = 0.86), as well as those studies in which all subjects were not taking any type of medication that could affect lipids and lipoproteins vs. those studies in which one or more subjects were taking drugs that might affect lipids and lipoproteins ( $Q_p = 2.9, p = 0.09$ ). Greater reductions in TC were associated with more recent year of publication (R = 0.41, 99% CI 0.07 to 0.66) and decreases in percent body fat (R = 0.47, 99% CI 0.01 to 0.76). No other statistically significant or clinically relevant associations were found between decreases in TC and other variables that were regressed, including baseline values of TC (R = 0.17, 99% CI -0.19 to 0.49).

**HDL-C**—Across all categories, a statistically significant increase of approximately 3% was observed. No statistically significant publication bias was observed (p = 0.25). With those groups in which one or more subjects were taking HRT deleted from the model, increases in HDL-C continued to be statistically significant ( $1.6 \pm 0.6$  mg/dl, 95% CI 0.2 to 3.3 mg/dl). In addition, changes in HDL-C remained statistically significant when we deleted the study in

which all subjects were hyperlipidemic  $(1.7 \pm 0.9 \text{ mg/dl}, 95\% \text{ CI} 0.2 \text{ to } 3.5 \text{ mg/dl})$  as well as those groups in which changes in diet occurred  $(1.9 \pm 0.9 \text{ mg/dl}, 95\% \text{ CI} 0.2 \text{ to } 3.8 \text{ mg/dl})$ . No statistically significant or clinically relevant differences were observed for those variables in which subgroup analyses were performed, including differences in HDL-C between premenopausal and postmenopausal women (Q<sub>b</sub> = 0.18, *p* = 0.67), as well as those studies in which all subjects were not taking any type of medication that could affect lipids and lipoproteins vs. those studies in which one or more subjects were taking drugs that might affect lipids and lipoproteins (Q<sub>b</sub> = 0.2, *p* = 0.70). A statistically significant and positive association was observed between HDL-C and V<sub>02</sub>max, with greater increases in HDL-C associated with greater increases in V<sub>02</sub>max (*R* = 0.50, 99% CI 0.10 to 0.75). No other statistically significant or clinically relevant associations were found between increases in HDL-C and other variables that were regressed, including baseline values of HDL-C (*R* = 0.31, 99% CI -0.38 to 0.59).

**LDL-C**—Across all categories, a statistically significant reduction of approximately 3% was observed for LDL-C. No statistically significant publication bias was observed (p = 0.46). With those groups in which one or more subjects were taking HRT deleted from the model, decreases in LDL-C continued to be statistically significant ( $-4.2 \pm 1.1 \text{ mg/dl}$ , 95% CI -6.4 to -2.0 mg/dl). In addition, changes in LDL-C remained statistically significant when we deleted the study in which all subjects were hyperlipidemic ( $-4.4 \pm 1.1 \text{ mg/dl}$ , 95% CI -6.5 to -2.2 mg/dl) as well as those groups in which changes in diet occurred ( $-4.3 \pm 1.1 \text{ mg/dl}$ , 95% CI -6.5 to -2.0 mg/dl). No statistically significant or clinically relevant differences were found for those variables in which subgroup or regression analyses were performed, including differences in LDL-C between premenopausal and postmenopausal women ( $Q_b = 0.32$ , p = 0.57), studies in which all subjects were not taking any type of medication that could affect lipids and lipoproteins vs. those studies in which one or more subjects were taking drugs that might affect lipids and lipoproteins ( $Q_b = 0.2$ , p = 0.67), and baseline levels of LDL-C (R = 0.12, 99% CI -0.26 to 0.47).

**TG**—Across all categories, a statistically significant decrease in TG of approximately 5% was observed. No statistically significant publication bias was observed (p = 0.31). With those groups in which one or more subjects were taking HRT deleted from the model, decreases in TG continued to be statistically significant ( $-3.6 \pm 2.0 \text{ mg/dl}$ , 95% CI -7.7 to -0.1 mg/dl). In addition, changes in TG remained statistically significant when we deleted the study in which all subjects were hyperlipidemic  $(-4.2 \pm 2.1 \text{ mg/dl}, 95\% \text{ CI} - 8.4 \text{ to} -0.1 \text{ mg/dl})$  as well as those groups in which changes in diet occurred ( $-4.2 \pm 2.0 \text{ mg/dl}$ , 95% CI -8.7 to -0.2 mg/dl). No statistically significant or clinically relevant differences were observed for any of the subgroup analyses performed. Greater decreases in TG were associated with more recent year of publication (R = 0.38, 99% CI 0.04 to 0.64) and lower study quality (R = 0.44, 99% CI 0.10 to 0.68). No other statistically significant or clinically relevant differences or associations were found for those variables in which subgroup or regression analyses were performed, including differences in TG between premenopausal and postmenopausal women ( $Q_{\rm b} = 1.28, p = 0.28$ ), those studies in which all subjects were not taking any type of medication that could affect lipids and lipoproteins vs. those studies in which one or more subjects were taking drugs that might affect lipids and lipoproteins ( $Q_b = 0.7$ , p = 0.42), and baseline values of TG (R = 0.28, 99% CI -0.07 to 0.57).

#### Secondary outcomes

Secondary outcomes are shown in Table 4. As can be seen, statistically significant exercise minus control group decreases of approximately 2% were found for both body weight and BMI, whereas an approximately 4% decrease was observed for percent body fat. A statistically significant exercise minus control group increase of approximately 13% was found for  $V_{02}max$ .

#### Discussion

The overall results of this study suggest that improvements of 2%-5% in lipids and lipoproteins can be expected as a result of an aerobic exercise program in women. However, although we found statistically significant improvements in TC, HDL-C, LDL-C, and TG, a moderate to large amount of heterogeneity was observed for all outcomes. For TC, some of this heterogeneity may be explained by the statistically significant associations we observed between decreases in TC with more recent year of publication and decreases in percent body fat. The association between decreases in TC with decreases in percent body fat suggests that some of the heterogeneity for changes in TC may be the result of differing changes in percent body fat across the different studies. In other words, greater decreases in body fat may have resulted in greater decreases in TC and, thus, greater heterogeneity in TC because of varying changes in percent body fat across the different studies. This leads us to question whether the decreases in TC were the result of the aerobic exercise program itself, the decreases in percent body fat that occurred as a result of the exercise program, or some combination of the two. Although the relationship between greater decreases in TC with decreases in percent body fat was not surprising, the relationship between greater decreases in TC with more recent year of publication was not expected. This latter relationship suggests that some other underlying factor(s) that we could not identify may be associated with changes in TC. Alternatively, our results could have been nothing more than a spurious finding, although we found the same relationship for TG.

Some of the heterogeneity in our HDL-C results may be partly explained by the association between increases in  $V_{02}max$  with increases in HDL-C. In other words, greater increases in  $V_{02}max$  may have resulted in greater increases in HDL-C and, thus, greater heterogeneity in HDL-C because of varying changes in HDL-C across the different studies. We were unable to identify any potential source or sources of heterogeneity for LDL-C, but it is important to note that the percentage of inconsistency for LDL-C was less than that of the other lipid and lipoprotein variables reported. Similar to changes in TC, the relationship between changes in TG and year of publication suggests that some other underlying factor(s) that we could not identify may be associated with changes in TG. In addition, lower-quality studies may have accounted for some of the heterogeneity in TG outcomes. However, the use of quality rating scales needs to be interpreted with extreme caution.<sup>45</sup>

The results of our meta-analysis differ some-what from the results of a previous meta-analysis published in 1989<sup>44</sup> dealing with the effects of exercise on lipids and lipoproteins in women. In this previous work, it was reported that exercise favorably altered TC and TG but not HDL-C or LDL-C. One possible reason for these discrepancies may have to do with differences in the inclusion criteria between the two studies. For example, in our current study, we limited our studies to randomized controlled trials, whereas the previous meta-analysis included randomized controlled trials, nonrandomized controlled trials, and uncontrolled trials. Another possible reason may have to do with different approaches used to analyze data in the two studies. For example, in our study we incorporated both between-study and within-study dispersion measures for estimating treatment effects, whereas the previous study limited its analysis to between-study dispersion only.

The most recent National Cholesterol Education Report on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults includes a recommendation of increased physical activity for improving lipids and lipoproteins.<sup>2</sup> As the vast majority of studies included in this meta-analysis appeared to follow the American College of Sports Medicine Guidelines for developing and maintaining cardiorespiratory fitness in adults,<sup>57</sup> in would seem appropriate to suggest that improvements of 2%–5% in lipids and lipoproteins could be expected by adhering to these guidelines. This includes aerobic exercise performed 3–5 days

per week for 20–60 continuous minutes at an intensity of 55%–90% of maximum heart rate, with lower intensity levels recommended for those who are more unfit. The use of aerobic exercise may be especially appropriate because numerous other benefits can be derived from it,<sup>58</sup> and the risk of injury, especially for walking, is generally low.<sup>59</sup> The former not withstanding, it is important to note that with the exception of LDL-C, the clinical importance of the changes for each of the lipids and lipoproteins observed in this study is not well known. For LDL-C, it has been suggested that for every 1% decrease, the relative risk for major coronary heart disease (CHD) events is reduced by approximately 1%.<sup>2</sup> As we found an approximate 3% decrease in LDL-C, we could expect an approximate 3% decrease in the relative risk for major CHD events. However, for those subjects in whom greater reductions are necessary, other types of lifestyle (e.g., diet) and pharmacological (e.g., statins) interventions may be necessary in conjunction with increased physical activity.<sup>2</sup>

Meta-analysis, like any other type of review, is limited by the available data. In fact, this is one of the major reasons for conducting a meta-analysis, that is, to use a quantitative approach for the purpose of providing directions for future research. For example, only one included study reported that all their female subjects were hyperlipidemic.<sup>30</sup> Because higher levels of lipids and lipoproteins increase one's risk for morbidity and mortality from CVD,<sup>2</sup> it would seem appropriate to suggest that future aerobic exercise training studies increase their focus on those women with elevated levels. Furthermore, as none of the included studies limited their inclusion criteria to women with CVD, future studies should consider such.

There was a lack of specific information about the timing of the menstrual cycle during the assessment of lipids and lipoproteins. This timing may be important because lipids and lipoproteins can vary depending on the phase of the menstrual cycle.<sup>60</sup> Therefore, it is suggested that future studies report, and editors publish, information about the timing of the menstrual cycle when assessing lipids and lipoproteins in women. Furthermore, it was sometimes difficult to discern if the women were taking any type of pharmacological intervention other than lipid-lowering agents that could affect lipids and lipoproteins (e.g., oral contraceptives, estrogen replacement therapy). These agents can affect lipids and lipoproteins, and it is suggested that future studies include, and editors publish, information about this. Finally, only two studies appeared to use the intention-to-treat approach in the analysis of their data.<sup>13</sup>,38 Thus, although aerobic exercise may be efficacious for reducing lipids and lipoproteins in women, we cannot reach any definitive conclusions about its effectiveness. Therefore, it would appear plausible to suggest that future studies include results using both intention-to-treat and analysis-by-protocol approaches.

In conclusion, the results of this study suggest that, overall, aerobic exercise is efficacious for improving TC, HDL-C, LDL-C, and TG in women.

#### Acknowledgements

This study was supported by a grant from the NIH National Heart, Lung and Blood Institute, award RO1-HL069802 (G.A. Kelley, Principal Investigator).

We thank William Haskell, Ph.D., Stanford University, for reviewing our reference list and providing suggestions for the coding of studies.

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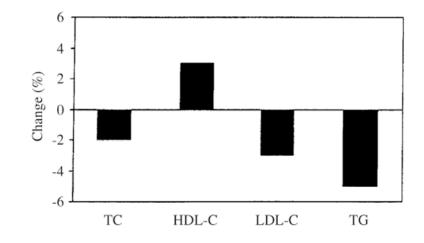
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Relative changes (%) in lipids and lipoproteins (exercise minus control). TC, total cholesterol; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; TG, triglycerides.

Study	Design	Subjects <sup>b</sup>	Exercise intervention	Lipids <sup>c</sup>	Lipid assessment
Agren et al. <sup>3</sup>	RCT <sup>e</sup>	38 women, included assignment to an exercise $(n = 23, \text{ age} = 22 \pm 3 \text{ years}^d)$ or control $(n = 15, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0, 0,$	14 weeks of aerobic activity, 3.5 days/week, at least 30 min/session, 70%–75% of Vo <sub>2</sub> max	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding eventice for 77 hours
Aldred et al. <sup>4</sup>	RCT	$agc = 21.0 \pm 1.0$ years) group 22 sedentary women, $41-55$ years of age, assigned to either an exercise ( $(n = 11)$ or control	12 weeks of walking, 2-7 days/week, 20-40 min/session, 63%-86% of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding
Allison et al. <sup>5</sup>	RCT	(n = 11) group 32 sedentary women, 17–26 years of age, assigned to either 30 $(n = 12)$ , or 45 $(n = 11)$ , min of eventias or to a control means $(n = 0)$	8 weeks of jogging, 3 days/week, 30 or 45 min/session, 85% MHR	TC, HDL, LDL, TG	exercise for 48 nours Morning after a 12-hour overninght fast and 24 hours after the lost evention section
Baker and Al- Naiadah <sup>6</sup>	RCT	or exercise, or to a control group $(n - j)$ 30 women assigned to either an exercise $(n = 20)$ or control $(n = 10)$ group	12 weeks of aerobic exercise, 2 days/week, 17 min/session, 65%–80% of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast
Busby et al. <sup>7</sup>	RCT	24 sedentary women 40–65 years of age assigned to an exercise group ( $n = 12$ , age 49.20 $\pm$ 18.01 years) or control group ( $n = 12$ , age = 49.90 $\pm$ 24.04 years)	12 weeks of walking/jogging, 3 days/week, 30 min/session, 70%–80% of MHR	TC, HDL, TG	Morning after a 10–12-hour overnight fast
Cauley et al. <sup>8</sup>	RCT	2004 $(n = 57.39 \pm 4.01)$ years) or control $(n = 104, a_{000} = 57.39 \pm 4.01)$ years) or control $(n = 104, a_{000} = 57.39 \pm 4.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 4.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$ years) or control $(n = 104, a_{000} = 57.39 \pm 1.17)$	104 weeks of walking, 3 days/week	TC, HDL, TG	Morning after fasting for at least 10 hours
Davison and Grant <sup>9</sup>	RCT	52 sedentary women, $40-60$ years of age assigned to either an exercise $(n = 48)$ or control $(n = 14)$	14 weeks of walking, 4 days/week, 30 min/ session, 70%–75% of MHR	TC, HDL, TG	NA
Duncan et al. <sup>10</sup>	RCT	53 sedentary women, 20–40 years of age, 53 sedentary women, 20–40 years of age, assigned to a strolling $(n = 18)$ , brisk walking $(n = 12)$ , aerobic walking $(n = 13)$ , or control $(n = 10)$ group	24 weeks of walking 4.8 km/day, 5 days/ week, at either 8 (aerobic walkers), 6.4 (brisk walkers), or 4.8 (strollers) km/h. at 86% (aerobic walkers), 67% (brisk walkers), or 56% (strollers) of MHR	TC, HDL, LDL, TG	Moming after a 12–14-hour overnight fast and avoiding exercise
Edin et al. <sup>11</sup>	RCT	17 sedentary women assigned to an exercise ( $n = 10$ ) or control ( $n = 7$ ) group	11 weeks of minitrampoline rebounding, 30 min/session, 5 days/week at 70%–85% of MHR	TC, HDL, TG	12+-hour fast
Fahlman et al. <sup>12</sup>	RCT	30 sedentary women assigned to an aerobic exercise $(n = 15, \text{ age } = 74 \pm 5 \text{ years})$ or control $(n = 15, \text{ are } = 74 \pm 5 \text{ years})$ or control	100000 1000 1000 1000 1000 1000 1000 1	TC, HDL, LDL, TG	Morning after a 12–14-hour overnight fast and avoiding exercise for 48–96 hours
Furukawa et al. 13	RCT	45 women, $32-57$ years of age, assigned to either an exercise $(n = 21)$ or control $(n = 24)$ group	12 weeks of walking, 2–3 days/week, 20–30 min/session, 65% of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast
Gillett et al. <sup>14</sup>	RCT	82 sedentary women, 49–59 years of age, assigned to either an exercise $(n = 23)$ or control $(n = 23)$ or num	16 weeks of low-impact dance exercise, 3.2 days/week, 28.5 min/session, 60%–80% of MHRR	TC, HDL	Morning after a 12-hour overnight fast
Grandjean et al. 15	RCT	37 sedentary women, 20 exercise and 17 control	24 weeks of walking/jogging/cycling, 3 days/ week, 20-60 min/session, 60%-80% of Vo.max	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast
Hagan et al. <sup>16</sup>	RCT	24 sedentary women (mean age = 35.2 years) assigned to either an exercise $(n = 12)$ or control $(n = 12)$ control	12 weeks of walking and/or running, 5 days/ week, 30 min/session	TC, HDL, LDL, TG	Morning after a 14-hour overnight fast
Hinkleman and Nieman <sup>17</sup>	RCT	(n - 12) group 36 sedentary women, 25–45 years of age, 18 exercise (age = 36.0 $\pm$ 6.8 years) and 18 control $(nn - 27.4 \pm 6.4 \text{ more})$	15 weeks of walking, 5 days/week, 45 min/ session, 62% of Vo <sub>2</sub> max	TC, LDL, TG	Morning after a 12-hour overnight fast and avoiding
Hopewell <sup>18</sup>	RCT	$dgc - 324 \pm 0.4$ years) 11 sedentary women, assigned to either an exercise $(n = 5)$ or control $(n = 6)$ group	24 weeks of walking, 3 days/week, 50 min/ session, 68% of Vo <sub>2</sub> max	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding evervise for at least 24 hour
Isler et al. <sup>19</sup>	RCT	45 sedentary women assigned to either a step aerobics $(n = 15, \text{ age} = 21.88 \pm 2.16 \text{ years})$ , aerobic dancing $(n = 15, \text{ age} 20.23 \pm 0.16 \text{ years})$ , or control $(n = 15, \text{ age} = 21.88 \pm 1.82 \text{ years})$ group	8 weeks of either step aerobics or aerobic dancing, 3 days/week, 20 min/session, 60%– 70% of MHRR	TC, HDL, LDL, TG	Morning after a 10–12-hour overnight fast

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Table 1

Characteristics of Included Studies<sup>a</sup>

Study	Design	$\mathrm{Subjects}^b$	Exercise intervention	Lipids <sup>c</sup>	Lipid assessment
Juneau et al. <sup>20</sup>	RCT	56 sedentary women, $40-60$ years of age, assigned to an exercise $(n = 28)$ or control $(n = 28)$ around	24 weeks of walking/jogging, 5 days/week, $47-54$ min/session, $50\%-60\%$ of $\mathrm{Vo}_2\mathrm{max}$	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast
Keller and Trevino <sup>21</sup>	RCT	35 second 35 sedentary women, 18–45 years of age, assigned to either 3× week exercise $(n = 12)$ , 5× week exercise $(n = 11)$ or control $(n = 12)$ from	24 weeks of walking, 3 or 5 days/week, 30 min/session, 50% of MHRR	TC, HDL, LDL, TG	Morning after a 7–11-hour overnight fast and after avoiding exercise
King et al. <sup>22</sup>	RCT	132 sedentary women, 50-65 years of age, assigned to high-intensity group-based exercise (n = 34), high-intensity home-based exercise $(n = 53)$ , hower-intensity home-based exercise $(n = 30)$ , or control $(n = 34)$ monu	52 weeks of walking/jogging/cycling, 3 days/ week, 40 min/session, 73%–88% of MHR (higher-intensity exercise groups) or 5 days/ week, 30 min/session, 60%–73% of MHR (housr-intensity exercise around)	HDL, LDL, TG	Moming after an overnight fast of at least 12 hours
Lee and White <sup>23</sup>	RCT	37 women, 40-61 years of age (mean age = 48.4 years), assigned to an exercise ( $n = 19$ ) or control ( $n = 18$ ) around	12 weeks of low-impact aerobics and walking, 3-4 days/week, 60% of MHRR	TC, HDL, TG	Morning after a 12-hour overnight fast
LeMura et al. <sup>24</sup>	RCT	22 sedentary women, assigned to either an aerobic exercise ( $n = 10$ , age = 21.0 ± 2.0 years) or control ( $n = 12$ are = 20.0 + 1.0 years) or control ( $n = 12$ are = 20.0 + 1.0 years)	16 weeks of either walking/jogging, cycling, or rowing, 3–4 days/week, 30–45 min/ session 70/8, -85%, of MHR	TC, HDL, LDL, TG	Morning after a 12–14-hour overnight fast and avoiding evervise for 24–48 hours
Lindheim et al. 25	RCT	.0 +1 %	24 weeks of walking/cycling, 3 days/week, 30 min/session, 70% of MHR	TC, HDL, LDL, TG	Morning after an overnight fast
Lovelady et al. <sup>26</sup>	RCT	33 women, assigned to either an exercise $(n = 18, 38 \text{ women}, 383\text{ second to either an exercise } (n = 18, 388 \text{ second to either an exercise} (n = 15, 388 \text{ second to either an exercise} 29.7 + 5.03 \text{ wears} around the either and around the either around $	12 weeks of walking, jogging, and cycling, 4.5 days/week, 20–45 min/session, 60%–70% of MHRR	TC, HDL, LDL, TG	Morning after fasting for at least 12 hours
Miller <sup>27</sup>	RCT	37 sedentary women assigned to either a 750 kcal/ week of exercise ( $n = 13$ ), 1500 kcal/week of	14 weeks of walking, 5 days/week, expending either 750 or 1500 kcal/week, 50% of Vo <sub>2</sub> max	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding
Nieman et al. <sup>28</sup>	RCT	exercise $(n = 14)$ , or control $(n = 10)$ group 43 sedentary women assigned to either an exercise $(n = 31)$ or control $(n = 22)$ aroun	12 weeks of walking, 5 days/week, 25–45 min/session 60%–80% of MHR	TC, HDL, LDL, TG	CACLOSCIOL 27 HOLES
Nieman et al. <sup>29</sup>	RCT	30 sedentary women, $67-85$ years of age, assigned to an exercise $(n = 14)$ or sham $(n = 16)$	12 weeks of walking, 5 days/week, 30–40 min/session, 60% of MHRR	TC, HDL, LDL, TG	NA
Ready et al <sup>30</sup>	RCT	25 postmenopausal women, 15 exercise and 10 25 postmenopausal women, 15 exercise and 10 control, $62 \pm 5.7$ years of age	24 weeks of walking, 5 days/week, 54 min/ session, 54% of MHRR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and 12 hours after the last everties essertion
Ready et al. <sup>31</sup>	RCT	53 sedentary women (mean age = 61.3 $\pm$ 5.8 years) assigned to either 3 days of exercise ( $n =$ 18, 5 days of exercise ( $n = 17$ ), or control ( $n = 18$ ) or on	24 weeks of walking, 3 days/week, 59 min/ session, 60% Vo <sub>2</sub> max, or 5 days/week, 56 min/session, 60% of Vo <sub>2</sub> max	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding exercise
Santiago et al. <sup>32</sup>	RCT	25 sedentary women, 20–40 years of age, 25 sedentary women, 20–40 years of age, assigned to either a walking $(n = 9, \text{ age} = 30.0 \pm 4.95 \text{ years})$ , jogging $(n = 8, \text{ age} = 28.75 \pm 5.12 \text{ years})$ , or control $(n = 8, \text{ age} = 29.38 \pm 5.71 \text{ years})$	20 weeks of either walking or jogging, 4 days/ week, 53 (walking) or 34 (jogging) min/ session, 73% (walking) or 84% (jogging) of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding exercise
Santiago et al. <sup>33</sup>	RCT	27 sedentary women, 22–40 years of age, assigned to either an exercise ( $n = 16$ , age = 30.1 $\pm 5.3$ years) or control ( $n = 11$ , age = 31.5 $\pm 6.1$ years) or control ( $n = 11$ , age = 31.5 $\pm 6.1$	40 weeks of walking, 4 days/week, 72% of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding exercise
Seals et al. <sup>34</sup>	RCT	years) group 35 sedentary women assigned to an exercise ( $n =$ 18, age = 62 ± 9 years) or no exercise and sodium restriction ( $n = 17$ , are = 65 ± 10, wears) or on n	13 weeks of walking, 5.8 days/week, 40 min/ session, 70% of MHR	TC, HDL, LDL, TG	Morning after an overnight fast
Sinatra et al. <sup>35</sup>	RCT	26 women, 28-64 years of age, assigned to an exercise $(n = 15)$ or control $(n = 11)$ group	12 weeks of walking, 3 times per week	TC, HDL, TG	Morning after an overnight fast, within 48 hours after the last exercise session
Stefanick et al. <sup>36</sup>	RCT	88 women, 45–64 years of age, assigned to either an exercise $(n = 43)$ or control $(n = 45)$ group	52 weeks of aerobic activity, 3 days/week, 60 mit/session	TC, HDL, LDL, TG	Moming after a 12–16-hour overnight fast and at least 12

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Study	Design	Subjects <sup>b</sup>	Exercise intervention	Lipids <sup>c</sup>	Lipid assessment
Sugiura et al. <sup>37</sup>	RCT	27 sedentary women, 40–60 years of age, assigned to an exercise $(n = 14, \text{ age} = 48.6 \pm 4.2 \text{ years})$ or control $(n = 13, \text{ age} = 48.0 \pm 3.6 \text{ years})$	104 weeks of walking/jogging/rhythmic dance, 1 or more days/week, 30 min/session 40%–60% of MHRR	TC, HDL, LDL, TG	hours after the last exercise session NA
Suter and Marti <sup>38</sup>	RCT	of age group 32 sedentary women, 25–55 years of age assigned to an exercise $(n = 16, age = 38.20 \pm 9.40$ years) or control $(n = 16 a a = 47, 70 + 8, 70)$ ears)	16 weeks of jogging, 2–6 days/week, 20–60 min/session (120 total min/week), 75%–87% of MHR	TC, HDL, LDL, TG	Morning after a 12–14-hour overnight fast
Thomas et al. <sup>39</sup>	RCT		12 weeks of running either 4 miles or 2 miles, 3 days/week, 30–36 (4-mile group) or 15–18 0-mile groun) min/session 75%, of MHR	TC, HDL, TG	Morning after a 12-hour overnight fast and 48 hours after the last eventies escion
Verity and Ismail <sup>40</sup>	RCT	10 sedentary women, 50–70 years of age, assigned to either an exercise $(n = 5, \text{ age} = 61.20 \pm 9.17)$ years) or control $(n = 5, \text{ age} = 57.20 \pm 8.27)$	16 weeks of walking, 3 days/week, 60–90 min/session, 65%–80% of MHRR	TC, HDL	Morning after a 10–12-hour overnight fast
Vetro <sup>41</sup>	RCT	Journal group 12 sedentary women, 24–59 years of age, assigned to either an exercise $(n = 6)$ or control (n = 6) or out	10 weeks of aerobic dance, 3 day/week, 20– 25 min/session, 70%–85% of MHR	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding evervice for at least 24 hours
Wallberg- Henriksson et al. 42	RCT	13 second assigned to either an exercise (her $36 \pm 36 \pm 4.9$ years) or control $(n = 7, \text{ age } = 35 \pm 5.3$ years) group	20 weeks of cycling, 5.2 days/week, 15 min/ session, 60%–90% of Vo <sub>2</sub> max	TC, HDL, LDL, TG	Morning after an overnight fast
Ziemer <sup>43</sup>	RCT	49 sedentary women, 23-49 years of age, assigned to either stair-stepper $(n = 7)$ , treadmill $(n = 13)$ , cross-country skier $(n = 13)$ , cycling $(n = 9)$ , or control $(n = 7)$ conditions	12 weeks of aerobic exercise using either a stair-stepper, treadmill, cross-country ski simulator, or stationary bicycle, 3 days/week, 20–45 min/session, at a self-selected intensity	TC, HDL, LDL, TG	Morning after a 12-hour overnight fast and avoiding exercise
<sup><i>a</i></sup> Description of studies limited to those subjects and <i>b</i> .	lies limited to t	$^{a}$ Description of studies limited to those subjects and variables that met our inclusion criteria.	là.		

Number of subjects limited to those in whom pre and post assessment of lipids took place.

 $^{c}$ Lipid variables listed limited to those that met our inclusion criteria, including availability.

dData reported as mean  $\pm$  SD.

e RCT, randomized controlled trial; kcal, kilocalories; MHRR, maximum heart rate reserve; Vo2max, maximum oxygen consumption; MHR, maximal heart rate; TC, total cholesterol; HDL-C, highdensity lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; TG, triglycerides; NA, not available.

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**Baseline Characteristics of Subjects** 

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		Exercise			Control	
Variable	na	$\mathbf{X} \pm \mathbf{SD}^{\hat{h}}$	Range <sup>c</sup>	u	$\mathbf{X} \pm \mathbf{SD}$	Range
Age, years	55	$41.6 \pm 14.3$	20.2–76.0	41	$42.9 \pm 14.4$	20.0-74.0
Height, cm	28	$163.5\pm2.8$	156.0 - 170.0	19	$163.3\pm3.7$	156.0-171.0
Body weight, kg	46	$67.8 \pm 8.1$	54.1 - 88.4	35	$68.2\pm8.9$	53.0-90.5
$BMI.^{d} (ke/m^{2})$	44	$25.5 \pm 3.4$	20.6–36.2	32	$25.6 \pm 3.8$	20.9-34.0
Body fat, %	33	$32.2 \pm 5.2$	21.7 - 43.1	24	$33.6 \pm 5.4$	21.3-43.4
$V_{0,max}$ , ml/kg <sup>-1</sup> /min <sup>-1</sup>	43	$28.8 \pm 6.2$	17.3-45.0	30	$27.6 \pm 6.4$	17.8-45.0
TC, mg/dl	53	$197.4\pm24.5$	159.7 - 257.9	41	$203.2\pm23.6$	155.0-264.0
HDL-C, mg/dl	56	$55.6\pm8.3$	35.2-80.8	42	$54.7\pm8.6$	31.2-69.2
LDL-C, mg/dl	46	$122.1 \pm 23.9$	79.8–178.6	33	$127.3 \pm 21.1$	78.2-187.5
TG, mg/dl	54	$100.9\pm27.3$	57.1–171.1	40	$103.1 \pm 31.1$	54.0-178.1

Kelley et al.

<sup>d</sup>BMI, body mass index; Vo2max, maximum oxygen consumption; TC, total cholesterol; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; TG, triglycerides.

 $^{\ensuremath{\mathcal{C}}}$  Range represents the means for each group from each study.

#### Table 3

### Training Program Characteristics

Variable	n <sup>a</sup>	$\mathbf{X} \pm \mathbf{SD}^{\boldsymbol{b}}$	Range
Length, weeks	56	$21.8 \pm 19.5$	8–104
Frequency, times/week	56	$3.7 \pm 1.1$	1–6
Intensity, %Vo <sub>2</sub> max	48	$69.2 \pm 10.1$	50-86
Duration, min/session	50	$36.3 \pm 13.2$	15-75
Compliance, % <sup>C</sup>	31	86.1 ± 13.5	31–52

 $a^{n}$ , total number of groups reporting data for that variable.

 $b_{\vec{X} \pm \text{SD}, \text{ mean } \pm \text{ standard deviation.}}$ 

<sup>c</sup>Compliance, percentage of exercise sessions attended.

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Variable	na	$\mathbf{X} \pm \mathbf{SD}^{\boldsymbol{b}}$	95% CI <sup>c</sup>	Q <sup>d</sup>	$p^{e}$	13
Primary outcomes, mg/dl						
$TC^{\vec{k}}$	53	$-4.3 \pm 1.3$	$-6.9 \text{ to } -1.7^{*}$	148.6	$<0.001^{**}$	65.
HDL-C	56	$1.8 \pm 0.9$	$0.1 \text{ to } 3.5^*$	500.6	$<0.001^{**}$	89
LDL-C	46	$-4.4 \pm 1.1$	$-6.5 \text{ to } -2.2^{*}$	108.3	$<0.001^{**}$	58
TG	53	$-4.2 \pm 2.1$	$-8.4 \text{ to } -0.1^{*}$	244.6	$<0.001^{**}$	78.7
Secondary outcomes						
Body weight, kg	39	$-1.0\pm0.2$	$-1.5 \text{ to } -0.6^{*}$	35.2	0.60	
BMI, kg/m <sup>2</sup>	18	$-0.6\pm0.2$	$-0.9 \text{ to } -0.3^{*}$	19.9	0.28	14.6
Body fat, %	29	$-1.3 \pm 0.3$	$-1.9 \text{ to } -0.7^{*}$	76.4	<0.001**	63
Vo <sub>2</sub> max, ml/kg <sup>-1</sup> /min <sup>-1</sup>	38	$3.5\pm0.4$	$2.7  ext{ to } 4.2^{*}$	302.6	$<0.001^{**}$	87.

 ${}^{D}X \pm \text{SEM}$ , mean  $\pm$  standard error of the mean.

 $^{c}$ CI, confidence interval.

<sup>d</sup>Q, heterogeneity value.

e p, significance value for Q.

 $f_{\rm I2},$  percentage (%) of inconsistency for study results, calculated from Q statistic.

<sup>g</sup>TC, total cholesterol; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; TG, triglycerides; BMI, body mass index; Vo2max, maximum oxygen consumption.

\* Significantly different from zero.

\*\* Statistically significant at p < 0.10.