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Association of Resistance Exercise, Independent of and Combined with Aerobic Exercise, with the Incidence of Metabolic Syndrome

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Abstract

Objective: To determine the association of resistance exercise, independent of and combined with aerobic exercise, with the risk of development of metabolic syndrome (MetS).

Patients and Methods: The study cohort included adults (mean ± SD age, 46±9.5 years) who received comprehensive medical examinations at the Cooper Clinic in Dallas, Texas, between January 1, 1987, and December, 31, 2006. Exercise was assessed by self-reported frequency and minutes per week of resistance and aerobic exercise and meeting the US Physical Activity Guidelines (resistance exercise ≥2 d/wk; aerobic exercise ≥500 metabolic equivalent min/wk) at baseline. The incidence of MetS was based on the National Cholesterol Education Program Adult Treatment Panel III criteria. We used Cox regression to generate hazard ratios (HRs) and 95% CIs.

Results: Among 7418 participants, 1147 (15%) had development of MetS during a median follow-up of 4 years (maximum, 19 years; minimum, 0.1 year). Meeting the resistance exercise guidelines was associated with a 17% lower risk of MetS (HR, 0.83; 95% CI, 0.73-0.96; P=.009) after adjusting for potential confounders and aerobic exercise. Further, less than 1 hour of weekly resistance exercise was associated with 29% lower risk of development of MetS (HR, 0.71; 95% CI, 0.56-0.89; P=.003) compared with no resistance exercise. However, larger amounts of resistance exercise did not provide further benefits. Individuals meeting both recommended resistance and aerobic exercise guidelines had a 25% lower risk of development of MetS (HR, 0.75; 95% CI, 0.63-0.89; P<.001) compared with meeting neither guideline.

Conclusion: Participating in resistance exercise, even less than 1 hour per week, was associated with a lower risk of development of MetS, independent of aerobic exercise.

Health professionals should recommend that patients perform resistance exercise along with aerobic exercise to reduce MetS.

List of abbreviations

BMI Body Mass Index

CI Confidence intervals

CVD Cardiovascular disease

HDL High-density lipoprotein cholesterol

HR Hazard ratios

MetS Metabolic Syndrome

PA Physical activity

SD Standard deviation

One third of US adults have metabolic syndrome (MetS)¹. Cardiometabolic disorders, such as glucose intolerance, insulin resistance, central obesity, dyslipidemia, and hypertension are its key components^{2, 3}. Therefore, MetS is an important risk factor for type 2 diabetes mellitus^{4, 5} and cardiovascular diseases (CVD)^{6, 7}. Increasing physical activity (PA) is a cornerstone for preventing and treating MetS^{3, 8}. Several intervention studies have shown the benefits of aerobic exercise for improving metabolic risk factors^{9, 10}.

Previous studies, mostly cross-sectional, have identified negative associations of muscular strength¹¹⁻¹⁴ or resistance exercise¹⁵⁻¹⁷ with the prevalence of MetS. Furthermore, recent cohort studies have indicated that higher levels of resistance exercise were associated with lower risks of type 2 diabetes mellitus in men and women¹⁸⁻²⁰, which suggests that increasing resistance exercise might be a potential target for preventing MetS. However, there is very little evidence from large epidemiological studies regarding the effects of resistance exercise on the development of MetS. Therefore, the aim of this study is to examine the association of resistance exercise, independent of/and combined with aerobic exercise, with the risk of developing MetS in relatively healthy middle-aged adults. We hypothesized that resistance exercise lowers the risk of developing MetS and the combination of resistance and aerobic exercise might be stronger associated with lower risk than either one independently.

METHODS

Study Population

The Aerobics Center Longitudinal Study is a cohort of men and women, who received extensive preventive medical examinations at the Cooper Clinic in Dallas, Texas during January 1st, 1987 and December 31st, 2006. Among 10 243 participants, we excluded 836

individuals with a history of myocardial infarction, stroke, or cancer and 1989 individuals with MetS at baseline. Our final sample included 7418 individuals (1384 women [19%]). The participants were predominantly non-Hispanic whites (>95%), well educated, and employed in, or retired from, professional or executive positions²¹. The Cooper Institute institutional review board annually approved the study, and written informed consents were obtained from participants before data collection at baseline and during follow-up examinations.

Clinical examination

All participants performed comprehensive medical examinations at baseline, including body composition assessments, blood chemistry analyses, blood pressure measurements, electrocardiography, physical examination, and detailed medical history questionnaire. Body mass index (BMI) was calculated from measured weight and height squared (kg/m²). Waist circumference was measured with anthropometric tape at the umbilicus level. Blood chemistry analyses, measuring triglycerides, high-density lipoprotein (HDL) cholesterol and fasting glucose, were obtained with automated bioassays after 12-hour fasting. Resting systolic and diastolic blood pressure were measured by standard auscultatory methods after 5 minutes of seated rest, and calculated as the average of at least two readings separated by 2 minutes.

Age, gender, smoking status, alcohol consumption, personal history of physician-diagnosed CVD, cancer, and parental history of CVD, hypertension, and diabetes were assessed by a medical history questionnaire. Heavy alcohol drinking was defined as >14 and >7 alcoholic drinks per week for men and women, respectively²². The medical history questionnaire included a PA questionnaire containing self-reported leisure-time PA or recreational PA during the past 3 months. We classified aerobic exercise into four categories: "inactive (0

MET-minutes/week)", "insufficient (1–499 MET-minutes/week)", "medium (500-999 MET-minutes/week)" and "high (≥1000 MET-minutes/week)" based on the 2008 US PA Guidelines²³.

Assessment of resistance exercise

Self-reported resistance exercise was assessed in the medical history questionnaire. Participants were asked about the weekly frequency and average exercise duration (minutes) for each session of muscle-strengthening PA using either free weights or weight training machines over the past 3 months. We used frequency $(0, 1, 2, 3, 4 \text{ or } \ge 5 \text{ times/week})$ and total amount $(0, 1-59, 60-119, 120-179 \text{ and } \ge 180 \text{ minutes/week})$ of resistance exercise, as well as meeting the 2008 PA Guidelines for resistance exercise ($\ge 2 \text{ times/week}^{23}$), as our main exposures. The total amount of resistance exercise was calculated by multiplying frequency of exercise with the average minutes per session.

Ascertainment of MetS

Participants were classified as having MetS using the criteria of the National Cholesterol Education Program Adult Treatment Panel III³ at both baseline and follow-up. MetS was based on the presence of 3 or more of the following risk factors: 1) abdominal or central obesity (waist circumference >102 cm in men, >88 cm in women), 2) fasting hypertriglyceridemia (≥150 mg/dL [to convert to mmol/L, multiply by 0.0113]), 3) low HDL cholesterol (<40 mg/dL in men, <50 mg/dL in women [to convert to mmol/L, multiply by 0.0259]), 4) high blood pressure (≥130/85 mm Hg or history of physician-diagnosed hypertension) and 5) high fasting glucose (≥100 mg/dL [to convert to mmol/L, multiply by 0.0555] or history of physician-diagnosed diabetes). Follow-up time was

calculated from the baseline examination to the first event of MetS or the last follow-up examination through 2006 for individuals who did not develop MetS.

Statistical Analysis

Baseline characteristics were summarized as mean and standard deviation (SD) for continuous variables, and as number and percentage (%) for categorical variables. Baseline differences for participants with different amounts of resistance exercise were examined using analyses of variance (ANOVA) for continuous variables and chi-squared tests for categorical variables.

Cox proportional hazard regression was used to compute hazard ratios (HRs) and their 95% confidence intervals (CIs) of MetS across different amounts and frequencies of resistance exercise. Participants who reported no resistance exercise were used as reference category. The regression models were adjusted for age, gender, examination year, BMI, current smoking, heavy alcohol drinking, abnormal electrocardiography, parental history of CVD, hypertension, diabetes, and aerobic exercise (inactive, insufficient, medium, and high). In addition, we examined the independent and combined effects of meeting aerobic (≥500 MET/week²³) and/or resistance exercise guidelines on the risk of developing MetS in the combined analyses.

To examine potential effect modification by sex in the association between resistance exercise and incident MetS, we tested interaction terms of sex and resistance exercise using Cox regression. In addition, we compared risk estimates in sex-stratified analyses. We did not find any significant interaction, and trends of developing MetS in men and women were similar. Therefore, we presented the results of pooled analyses. All statistical tests were 2-

sided, and significance was set at *P*<.05. All analyses were conducted using SAS software, version 9.4.

RESULTS

Among 7418 participants, 15% (n=1147) developed MetS during a median follow-up of 4 years (maximum 19 years; minimum, 0.1 year) (Table 1). Among individuals who participated in resistance exercise (n=2785 [38%]), resistance exercise was most frequently performed for 60-119 minutes per week (n=1061 [38%]). Compared to individuals not performing resistance exercise, individuals with higher levels of resistance exercise were more likely to be younger, leaner (lower BMI and waist circumference), and aerobically active. However, the proportion of men decreased with higher levels of resistance exercise. Individuals who participated in resistance exercise were also less likely to smoke and had more favorable lipids profile (lower triglycerides and higher HDL cholesterol; all *P*<.05).

Performing any resistance exercise was associated with a 17% lower risk of developing MetS (HR, 0.83; 95% CI, 0.72-0.95; P=.006) after adjusting for potential confounders, including aerobic exercise levels in the fully adjusted model 3 (Table 2). Meeting the resistance exercise guidelines had a similar 17% lower risk of MetS (HR, 0.83; 95% CI, 0.73-0.96; P=.009) in the full model (model 3). Furthermore, we found that resistance exercise at 1-59, 60-119, 120-179, and \geq 180 minutes per week were all associated with lower HRs for MetS (all P<.05), compared to no resistance exercise; after adjusting for age, gender, and examination year (model 1). However, after further adjustment for other potential confounders and aerobic exercise levels (model 3), only 1-59 minutes per week of resistance exercise was associated with a 29% reduced risk of MetS (HR, 0.71; 95%CI, 0.56-0.89; P=.003). We also found that

four days per week of resistance exercise was associated with a 38% lower risk of developing MetS (HR, 0.62; 95%CI, 0.44-0.89; P=.009), compared to no resistance exercise in the fully adjusted model (model 3). In additional analyses after further adjustment for the number of MetS risk factors (0, 1, or 2) at baseline, the results were virtually the same, in that the risk of developing MetS was 14% lower in individuals performing any resistance exercise (HR, 0.86; 95% CI, 0.75-0.98; P=.02), 14% lower in individuals meeting the recommended guidelines (HR, 0.86; 95% CI, 0.75-0.99; P=.03), 26% lower in individuals performing <1 hour per week (HR, 0.74; 95% CI, 0.58-0.93; P=.01), and 33% lower in individuals performing 4 times per week (HR, 0.67; 95% CI, 0.47-0.95; P=.03) resistance exercise.

In addition, we examined the risk of MetS among individuals with the same total amount of weekly resistance exercise (minutes/week), but at different frequencies (1-2 vs \geq 3 times/week). For example, some people may perform 2 hours of weekly resistance exercise in one or two sessions, especially during weekends (so-called "weekend warriors"), whereas others may perform the same 2 hours of weekly resistance exercise in more than 2 sessions.

The joint analysis of frequency and the total amount of resistance exercise (Figure 1) did not show any significant differences in the risk of developing MetS between less frequent (1-2 times/week) and more frequent (≥3 times/week) exercisers among individuals with the same total amount of weekly resistance exercise (all P>.05). However, we observed a 33% lower risk of developing MetS (HR, 0.67; 95%CI, 0.49-0.91; *P*=.01) in individuals who performed resistance exercise 1-2 times per week with a total exercise amount of 1-59 minutes per week. Further, we found no difference in incident MetS in individuals

performing 1 to 59 minutes per week of resistance exercise for less than 1 year and more than 1 year (P>.05).

Figure 2 illustrates the independent and combined associations of meeting the resistance and/or aerobic exercise guidelines with incident MetS. We found that individuals meeting both recommended resistance and aerobic exercise guidelines had a 25% lower risk of developing MetS (HR, 0.75; 95% CI, 0.63-0. 89; *P*<.001), compared to individuals meeting neither guidelines.

DISCUSSION

This large cohort study yielded 3 major study findings. First, we demonstrated that participating in resistance exercise, independent of aerobic exercise, significantly decreases the risk of developing MetS, compared to no resistance exercise in a middle-aged relatively healthy population (P=.006). Specifically, less than one hour per week of resistance exercise resulted in significantly lower risk of MetS compared to no resistance exercise (P=.003). However, higher volumes of resistance exercise did not provide further benefits (Table 2), suggesting against the "more is better" philosophy. Second, the combined analysis of weekly frequency and total amount of resistance exercise (Figure 1) showed no effect of exercise frequency in incident MetS at a given total volume of resistance exercise. Therefore, resistance exercise for less than one hour per week, regardless of training frequency, may be important in preventing MetS. Third, meeting both resistance and aerobic exercise guidelines was associated with 25% lower risk of developing MetS, compared to meeting neither of these guidelines (Figure 2). This suggests additional benefits of doing both resistance and aerobic exercise for the prevention of MetS.

Previous studies have indicated a negative association of muscular strength and MetS, which was still present after adjusting for aerobic fitness¹². However, the protective effect of muscular strength against MetS might be explained by regular participation in resistance exercise, because resistance exercise is a major determinant of muscular strength^{24, 25}.

Cross-sectional studies of muscle-strengthening PA have also reported a negative association with the prevalence of MetS¹⁵⁻¹⁷, which is in line with our findings. Nevertheless, those prior studies only investigated the effect of participating in resistance exercise (yes/no) or meeting the resistance exercise guidelines (yes/no). On the other hand, our study further examined the dose-response relationship between resistance exercise and incident MetS across different weekly frequencies and total amounts of resistance exercise. In addition, we also examined the independent and combined effects of resistance and aerobic exercise on the development of MetS.

Several studies have investigated the associations between resistance exercise and type 2 diabetes mellitus, another common metabolic disease. Grøntved et al. found a reduced risk of type 2 diabetes mellitus by performing less than one hour of resistance exercise per week in 32 000 men and 99 000 women ^{19, 20}. In addition, they showed that a combination of aerobic and resistance exercise was superior in preventing type 2 diabetes mellitus. We found similar results for the prevention of MetS. Further, they found a linear dose-response relationship between the amount of resistance exercise and the risk of incident type 2 diabetes mellitus. In contrast, however, we did not observe a linear dose-response relationship between resistance exercise and the risk of developing MetS, suggesting against the "more is better" hypothesis regarding resistance exercise and development of MetS.

However, this might be at least partially due to the smaller sample size and number of cases in our study. It is also possible that resistance exercise dose-response curves may be different between MetS and type 2 diabetes mellitus. These contradictory findings suggest that further investigations on dose-response relationships between resistance exercise and different health outcomes are clearly warranted. We also investigated the dose-response relationship between the frequency of resistance exercise and risk of MetS, demonstrating significant benefits of four times per week resistance exercise. However, this result is somewhat complicated since the frequency does not necessarily fully represent the total amount of resistance exercise. Therefore, the prescription of frequency in the current resistance exercise guidelines may lack sufficient detail, whereas a prescription of total minutes per week might be more appropriate.

The current study demonstrated that there is no significant difference in the risk of MetS between 1-59 and ≥180 minutes per week of resistance exercise, which suggests no additional benefits of higher levels of resistance exercise on the development MetS. In addition, the dose-response relationship between resistance exercise and MetS may not be linear, but reverse J-shaped, which has been found in studies regarding aerobic exercise and CVD health²⁶⁻²⁸. Although it is not clear why there are no further benefits on incident MetS by increasing the amount of resistance exercise, it may be related to no significant differences in blood pressure and fasting glucose across different amounts of resistance exercise, as shown in Table 1. However, more favorable lipid profiles (Triglycerides and HDL cholesterol) by increasing resistance exercise (Table 1) may partially explain the benefits of resistance exercise on the development of MetS since blood lipids are the components of MetS. Furthermore, additional analyses did not show significant differences in risk of MetS in

individuals performing weekly 1-59 minutes resistance exercise for less than one year and more than one year (*P*>.05). A possible explanation could be the absence of training progression (no gradual increase in amount and/or intensity of resistance exercise) after a certain period, which results in a stabilization of the muscle mass and strength, and therefore no further health benefits. Future studies of long-term resistance exercise training with different doses and intensities are therefore needed to determine the protection against MetS as well as CVD.

In 2004, Lee et al.²⁹ introduced the concept of 'weekend warriors', individuals who meet the aerobic exercise guidelines but performed their PA in 1-2 days per week, possibly during weekends. They demonstrated that 'weekend warriors' still had mortality benefits, compared to sedentary individuals, but their benefits were less, compared to individuals who were regularly physically active, especially in individuals with major CVD risk factors, such as smoking, overweight, and hypertension. In our study there was no effect of increased frequency with the same amount of resistance exercise (all *P*>.05). Nevertheless, only individuals performing 1-59 minutes of resistance exercise in 1-2 sessions per week had significantly lower risk of MetS, compared to no resistance exercise (P=.01). This suggests that even a relatively small amount of resistance exercise once or twice per week may be enough to maximally reduce the risk of MetS, at least from the resistance exercise perspective. However, it should be mentioned that the sample sizes and number of cases were smaller in categories with higher levels of resistance exercise, which reduced the statistical power in these groups.

MetS is more prevalent in older and overweight individuals¹. However, subgroup analyses in our study appear to show similar negative trends, although not significant, for resistance exercise and MetS in different BMI (<25 vs ≥25 kg/m²) and age (<50 vs ≥50 years old) groups (data not shown). The lack of statistically significance was probably due to the small number of participants and MetS cases across these strata. Nevertheless, the reduced risk of MetS by resistance exercise remained significant after adjusting for BMI and age, and shows consistency in our findings.

The strengths of this study include a large cohort with a relatively long follow-up time. Furthermore, we believe that this is the first prospective study that investigated the association between resistance exercise and incident MetS. However, limitations of our study include self-reported data on PA, which may cause measurement errors due to overreporting of leisure-time PA³⁰. Nevertheless, over-reporting generally causes an underestimation of the true effect of exercise on health outcomes³¹. Only baseline levels of PA were used for the analyses, therefore changes in PA patterns were not included in the study. Our study includes primarily well-educated non-Hispanic whites from middle-to-upper socioeconomic strata, which may limit the generalizability of the results, thus the findings may be different in other populations. Conversely, homogeneity in ethnicity and socioeconomic status reduces potential confounding by race/ethnicity, education, and income. Physiological characteristics of this cohort are also similar to other representative population samples²¹. Another limitation is that we had no information about medications to take into account in the analyses. Although we adjusted for potential confounders such as medical conditions (e.g., hypertension, diabetes, and abnormal electrocardiography) and lifestyle

factors (e.g., smoking, alcohol intake, and body mass index), randomized controlled trials of resistance exercise are warranted to remove those confounding biases in the future.

CONCLUSION

Meeting the resistance exercise guidelines, independent of aerobic exercise, decreases the risk of developing MetS in a middle-aged adult population. Especially, relatively smaller amounts of resistance exercise, less than one hour in 1-2 sessions per week as could be seen in the "weekend warrior" profile, resulted in the highest reduction in the risk of developing MetS, compared to no resistance exercise. Also, meeting both resistance and aerobic exercise guidelines is superior in preventing MetS. Therefore, resistance exercise, independent of/and combined with aerobic exercise, should be included in one's PA routine for the prevention of MetS. Clinicians should routinely recommend resistance exercise training, in addition to aerobic training, for the prevention of MetS and future CVD risk. In addition, individuals with CVD risk factors should consider more individualized, safe and effective exercise program under the direction of a qualified exercise professional.

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References

- 1. Ford ES, Li C, Zhao G. Prevalence and correlates of metabolic syndrome based on a harmonious definition among adults in the US. J Diabetes. 2010;2(3):180-93.
- Eckel RH, Grundy SM, Zimmet PZ. The metabolic syndrome. Lancet.
 2005;365(9468):1415-28.
- National Cholesterol Education Program Expert Panel on Detection E, Treatment of High Blood Cholesterol in A. Third Report of the National Cholesterol Education
 Program (NCEP) Expert Panel on Detection, Evaluation, and Treatment of High Blood
 Cholesterol in Adults (Adult Treatment Panel III) final report. Circulation.
 2002;106(25):3143-421.
- 4. DeFina LF, Vega GL, Leonard D, Grundy SM. Fasting glucose, obesity, and metabolic syndrome as predictors of type 2 diabetes: the Cooper Center Longitudinal Study. J Investig Med. 2012;60(8):1164-8.
- 5. Ford ES, Li C, Sattar N. Metabolic syndrome and incident diabetes: current state of the evidence. Diabetes Care. 2008;31(9):1898-904.
- 6. Gami AS, Witt BJ, Howard DE, et al. Metabolic syndrome and risk of incident cardiovascular events and death: a systematic review and meta-analysis of longitudinal studies. J Am Coll Cardiol. 2007;49(4):403-14.
- 7. Mottillo S, Filion KB, Genest J, et al. The metabolic syndrome and cardiovascular risk a systematic review and meta-analysis. J Am Coll Cardiol. 2010;56(14):1113-32.
- 8. Lavie CJ, Milani RV. Cardiac rehabilitation and exercise training programs in metabolic syndrome and diabetes. J Cardiopulm Rehabil. 2005;25(2):59-66.

- Pattyn N, Cornelissen VA, Eshghi SR, Vanhees L. The effect of exercise on the cardiovascular risk factors constituting the metabolic syndrome: a meta-analysis of controlled trials. Sports Med. 2013;43(2):121-33.
- 10. Bateman LA, Slentz CA, Willis LH, et al. Comparison of aerobic versus resistance exercise training effects on metabolic syndrome (from the Studies of a Targeted Risk Reduction Intervention Through Defined Exercise STRRIDE-AT/RT). Am J Cardiol. 2011;108(6):838-44.
- 11. Wijndaele K, Duvigneaud N, Matton L, et al. Muscular strength, aerobic fitness, and metabolic syndrome risk in Flemish adults. Med Sci Sports Exerc. 2007;39(2):233-40.
- 12. Jurca R, Lamonte MJ, Church TS, et al. Associations of muscle strength and fitness with metabolic syndrome in men. Med Sci Sports Exerc. 2004;36(8):1301-7.
- 13. Jurca R, Lamonte MJ, Barlow CE, et al. Association of muscular strength with incidence of metabolic syndrome in men. Med Sci Sports Exerc. 2005;37(11):1849-55.
- 14. Artero EG, Lee DC, Lavie CJ, et al. Effects of muscular strength on cardiovascular risk factors and prognosis. J Cardiopulm Rehabil Prev. 2012;32(6):351-8.
- Magyari PM, Churilla JR. Association between lifting weights and metabolic syndrome among U.S. Adults: 1999-2004 National Health and Nutrition Examination Survey. J Strength Cond Res. 2012;26(11):3113-7.
- 16. Churilla JR, Magyari PM, Ford ES, Fitzhugh EC, Johnson TM. Muscular strengthening activity patterns and metabolic health risk among US adults. J Diabetes. 2012;4(1):77-84.
- 17. Churilla JR, Johnson TM, Magyari PM, Crouter SE. Descriptive analysis of resistance exercise and metabolic syndrome. Diabetes Metab Syndr. 2012;6(1):42-7.

- Crump C, Sundquist J, Winkleby MA, Sieh W, Sundquist K. Physical Fitness Among Swedish Military Conscripts and Long-Term Risk for Type 2 Diabetes Mellitus: A Cohort Study. Ann Intern Med. 2016.
- 19. Grontved A, Rimm EB, Willett WC, Andersen LB, Hu FB. A prospective study of weight training and risk of type 2 diabetes mellitus in men. Arch Intern Med. 2012;172(17):1306-12.
- 20. Grontved A, Pan A, Mekary RA, et al. Muscle-strengthening and conditioning activities and risk of type 2 diabetes: a prospective study in two cohorts of US women. PLoS Med. 2014;11(1):e1001587.
- 21. Blair SN, Kannel WB, Kohl HW, Goodyear N, Wilson PW. Surrogate measures of physical activity and physical fitness. Evidence for sedentary traits of resting tachycardia, obesity, and low vital capacity. Am J Epidemiol. 1989;129(6):1145-56.
- 22. National Institute on Alcohol Abuse and Alcoholism. Alcohol use and alcohol use disorders in the United States: Main findings from the 2001-2002 National Epidemiologic survey on alcohol and related conditions (NESARC). NIH Publication No. 05-5737 2006.
- 23. US Department of Health and Human Services. Physical Activity Guidelines for Americans. Washington (DC): US Department of Health and Human Services; 2008. http://www.health.gov/PAGuidelines. Accessed December 2, 2016
- 24. Thomis MA, Beunen GP, Maes HH, et al. Strength training: importance of genetic factors. Med Sci Sports Exerc. 1998;30(5):724-31.
- 25. Williams MA, Haskell WL, Ades PA, et al. Resistance exercise in individuals with and without cardiovascular disease: 2007 update: a scientific statement from the

- American Heart Association Council on Clinical Cardiology and Council on Nutrition, Physical Activity, and Metabolism. Circulation. 2007;116(5):572-84.
- 26. Eijsvogels TM, Molossi S, Lee DC, Emery MS, Thompson PD. Exercise at the Extremes:

 The Amount of Exercise to Reduce Cardiovascular Events. J Am Coll Cardiol.

 2016;67(3):316-29.
- 27. Lavie CJ, O'Keefe JH, Sallis RE. Exercise and the heart--the harm of too little and too much. Curr Sports Med Rep. 2015;14(2):104-9.
- 28. Lavie CJ, Arena R, Swift DL, et al. Exercise and the cardiovascular system: clinical science and cardiovascular outcomes. Circ Res. 2015;117(2):207-19.
- 29. Lee IM, Sesso HD, Oguma Y, Paffenbarger RS, Jr. The "weekend warrior" and risk of mortality. Am J Epidemiol. 2004;160(7):636-41.
- 30. Adams SA, Matthews CE, Ebbeling CB, et al. The effect of social desirability and social approval on self-reports of physical activity. Am J Epidemiol. 2005;161(4):389-98.
- 31. Celis-Morales CA, Perez-Bravo F, Ibanez L, et al. Objective vs. self-reported physical activity and sedentary time: effects of measurement method on relationships with risk biomarkers. PLoS One. 2012;7(5):e36345.

Figures

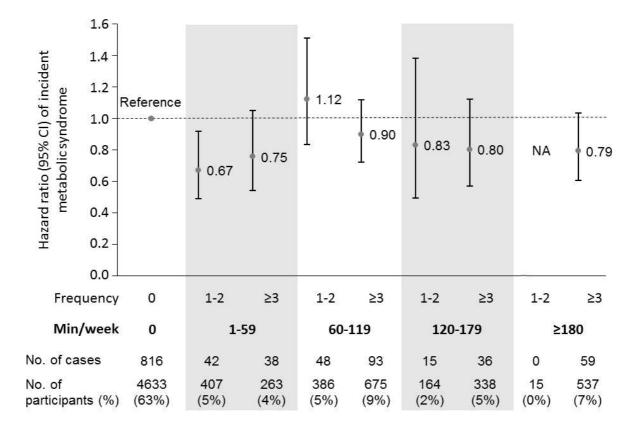


Figure 1. Hazard ratios for metabolic syndrome by the combination of weekly frequency (1-2 vs——3 times/wk) and minutes of resistance exercise (0, 1-59, 60-119, 120-179 and ≥180 min/wk). The dots indicate hazard ratios and the lines present 95% Cls. The model was adjusted for age (years), sex, examination year, body mass index (kg/m²), current smoking (yes/no), heavy alcohol drinking (yes/no), abnormal electrocardiographic findings (yes/no), parental history of cardiovascular disease, hypertension, and diabetes (yes/no for each), and aerobic exercise (inactive, insufficient, medium, and high). Analysis in the category of 180 minutes or more in 1 to 2 sessions of resistance exercise per week was not applicable (NA).

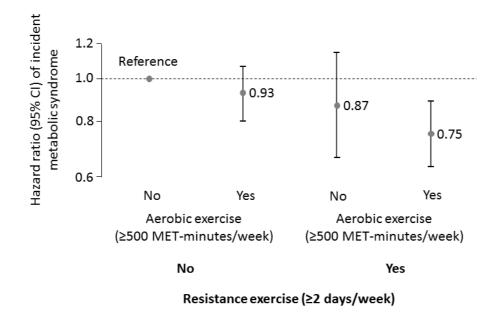


Figure 2. Hazard ratios for metabolic syndrome by meeting the 2008 US Physical Activity

Guidelines for resistance (≥2 d/wk) and aerobic (≥500 metabolic equivalent [MET] min/wk)

activities at baseline. The dots present hazard ratios and the lines the 95% CIs. The model

was adjusted for age (years), sex, examination year, body mass index (kg/m²), current

smoking (yes/no), heavy alcohol drinking (yes/no), abnormal electrocardiographic findings

(yes/no), and parental history of cardiovascular disease, hypertension, and diabetes (yes/no for each).

Table 1. Baseline characteristics of the 7418 participants by weekly minutes of resistance exercise^{a, b,}
c.

Characteristics	Weekly mir	Weekly minutes of resistance exercise (min/week)				
	0	1-59	60-119	120-179	≥180	
	(n=4633)	(n=670)	(n=1061)	(n=502)	(n=552)	
Age	46.7 (9.7)	45.9 (8.3)	46.2 (9.0)	45.1 (9.5)	43.7 (10.1)	<.001
Sex (male)	3795	568 (85%)	856 (81%)	369 (74%)	446 (81%)	<.001
	(82%)					
BMI (kg/m²)	25.3 (3.2)	24.9 (2.9)	24.9 (3.0)	24.8 (3.2)	24.8 (3.1)	<.001
Current smokers	522 (11%)	56 (8%)	93 (9%)	57 (11%)	60 (11%)	.04
Heavy alcohol drinking	562 (12%)	77 (11%)	132 (12%)	60 (12%)	64 (12%)	.98
Aerobic exercise						
(MET-min/week)						<.001
0	1125	40 (6%)	45 (4%)	32 (6%)	42 (8%)	
1-499	(24%)	89 (13%)	117 (11%)	65 (13%)	73 (13%)	
500-999	708 (15%)	135 (20%)	246 (23%)	115 (23%)	95 (17%)	
≥ 1000	899 (19%)	406 (61%)	653 (62%)	290 (58%)	342 (62%)	
	1901					
	(41%)					
Abnormal ECG	387 (8%)	53 (8%)	69 (7%)	29 (6%)	34 (6%)	.05
Parental history of	1177	162 (24%)	255 (24%)	130 (26%)	132 (24%)	.79
cardiovascular disease	(25%)					

Parental history of	1602	251 (37%)	379 (36%)	191 (38%)	184 (33%)	.28
hypertension	(35%)					
Parental history of	632 (14%)	91 (14%)	134 (13%)	61 (12%)	67 (12%)	.71
diabetes						
Metabolic syndrome						
Waist circumference	88.5 (11.0)	86.8 (10.3)	86.0 (10.7)	84.5 (11.1)	84.9 (10.3)	<.001
(cm)						
Triglycerides (mg/dL)	103.4	92.9 (44.3)	97.0 (52.4)	96.0 (53.1)	94.4 (59.6)	<.001
	(59.6)					
HDL cholesterol (mg/dL)	53.0 (14.5)	55.0 (14.6)	55.1 (14.4)	56.6 (14.7)	54.9 (14.3)	<.001
Systolic blood pressure	119 (13)	119 (13)	119 (13)	119 (13)	120 (13)	.67
(mm Hg)						
Diastolic blood pressure	79 (9)	80 (9)	79 (9)	79 (10)	80 (9)	.62
(mm Hg)						
Fasting glucose (mg/dL)	96.2 (10.1)	96.5 (13.0)	96.2 (11.9)	95.6 (11.4)	96.2 (13.3)	.77

^aBMI = body mass index; ECG = electrocardiographic findings; HDL-C = high-density lipoprotein cholesterol; MET = metabolic equivalent.

^bData are presented as mean ± SD or No. (percentage) of participants.

^cSI conversion factors: To convert triglycerides to mmol/L, multiply by 0.0113; to convert HDL-C to mmol/L, multiply by 0.0259; to convert glucose to mmol/L, multiply by 0.0555.

Table 2. Hazard Ratios for Metabolic Syndrome in 7418 Study Participants Stratified by Weekly Frequency and Minutes of Resistance Exercise^a.

	N (%)	No. of	Adjusted Hazard Ratio (95% CI)					
		cases	Model 1 ^b	Model 2 ^c	Model 3 ^d			
Weekly minutes of	Weekly minutes of resistance exercise (min/week)							
0	4633 (62%)	816	1.00 [Reference]	1.00 [Reference]	1.00 [Reference]			
1-59	670 (9%)	80	0.62 (0.49-0.78)	0.69 (0.55-0.86)	0.71 (0.56-0.89)			
60-119	1061 (14%)	141	0.83 (0.69-0.99)	0.93 (0.77-1.11)	0.96 (0.80-1.16)			
120-179	502 (7%)	51	0.66 (0.50-0.88)	0.78 (0.59-1.04)	0.81 (0.61-1.07)			
≥180	552 (7%)	59	0.65 (0.50-0.85)	0.76 (0.58-0.99)	0.78 (0.60-1.02)			
P-trend			<.001	.006	.03			
Any resistance exer	cise							
No (0 min/week)	4633 (62%)	816	1.00 [Reference]	1.00 [Reference]	1.00 [Reference]			
Yes (≥1 min/week)	2785 (38%)	331	0.71 (0.62-0.80)	0.80 (0.71-0.91)	0.83 (0.72-0.95)			
Weekly frequency o	Weekly frequency of resistance exercise (frequency/week)							
0	4633 (62%)	816	1.00 [Reference]	1.00 [Reference]	1.00 [Reference]			
1	206 (3%)	22	0.80 (0.53-1.23)	0.81 (0.53-1.24)	0.83 (0.54-1.27)			
2	766 (10%)	83	0.72 (0.58-0.91)	0.81 (0.65-1.02)	0.84 (0.67-1.06)			
3	1221 (16%)	163	0.77 (0.65-0.91)	0.86 (0.72-1.01)	0.88 (0.74-1.05)			
4	339 (5%)	32	0.48 (0.34-0.68)	0.60 (0.42-0.86)	0.62 (0.44-0.89)			
≥5	253 (3%)	31	0.66 (0.46-0.95)	0.79 (0.55-1.13)	0.81 (0.56-1.16)			
P-trend			<.001	.001	.005			

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No (<2 days/week)	4839 (65%)	838	1.00 [Reference]	1.00 [Reference]	1.00 [Reference]
Yes (≥2 days/week)	2579 (35%)	309	0.71 (0.62-0.80)	0.81 (0.71-0.92)	0.83 (0.73-0.96)

^a MetS = metabolic syndrome.

^b Adjusted for age, gender and examination year.

^c Adjusted for model 1 plus body mass index, current smoking, heavy alcohol drinking, abnormal electrocardiography, parental history of cardiovascular disease, hypertension, and diabetes.

^d Adjusted for model 2 plus aerobic exercise (inactive, insufficient, medium and high).