Effects of 24-Week Aerobic and Resistance Training on Carotid Artery Intima-Media Thickness and Flow Velocity in Elderly Women with Sarcopenic Obesity

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Aim: Sarcopenic obesity (SO) is closely associated with cardiovascular disease (CVD) in elderly women. Increases in body fat and decreases in muscle mass are closely associated with increased carotid intima-media thickness (CIMT). The aim of this study was to examine the influence of a 24-week aerobic and resistance training program on carotid parameters in SO.

Methods: Fifty elderly women (74.1 ± 6.1 years) with SO were randomly divided into an exercise group and a control group. The exercise group performed combined exercise over 24 weeks, consisting of resistance and aerobic training for 50-80 min, 5 times a week. Carotid variables were measured using B-mode ultrasound. The differences in the carotid variables and the relative changes between baseline and after 24 weeks were evaluated.

Results: In the analysis of variance (ANOVA) results, CIMT (p=0.013), systolic flow velocity (p=0.007), diastolic flow velocity (p=0.006), and wall shear rate (p=0.010) showed significant interactions. In paired t-test results of the exercise group, CIMT significantly decreased (p<0.01) and systolic flow velocity (p<0.01), diastolic flow velocity (p<0.001), and wall shear rate (p<0.05) significantly increased after 24 weeks.

Conclusion: The 24-week combined exercise effectively decreased CIMT and increased carotid flow velocity and wall shear ratio. Therefore, combined exercise is thought to contribute to the improvement of the risk of CVD in elderly women with SO.

Key words: Sarcopenic obesity, Combined exercise, Carotid intima-media thickness, Carotid flow velocity

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Introduction

A combination of sarcopenia and obesity is defined as sarcopenic obesity (SO)¹, ². SO is directly related to increased body fat mass (BFM) and decreased appendicular skeletal muscle mass (ASM) in the elderly¹, ². SO is associated with traditional cardiovascular disease (CVD) risk factors and mortality in the elderly¹, ³, ⁴. Carotid artery parameters such as carotid artery intima-media thickness (CIMT) and carotid artery flow velocity (FV) are reported to be independent risk factors for CVD⁵, ⁶. Increased BFM and decreased ASM are closely associated with increased CIMT⁷, ⁸. Moreover, increased BFM is associated with decreased FV as well as CIMT in the elderly⁷, ⁹. Some cross-sectional studies reported that SO is associated with reductions in physical activity, muscle
Aerobic exercise is also reported to be effective in improving CVD risk factors, including BFM and cardiorespiratory endurance. Furthermore, combined exercise (complex resistance and aerobic exercise) has the effect of decreasing BFM and increasing ASM, as well as improving CVD risk factors.

Previous evidence for exercise and CIMT suggests that exercise training does not affect CIMT in healthy young adults. However, exercise training has numerous beneficial effects in patients with either type 2 diabetes or CAD, and exercise plays an important role in the management of type 2 diabetes and in cardiac rehabilitation. It remains unclear whether regular exercise in elderly women positively modulates CIMT. In addition, the effect of a combined exercise program on CIMT in SO elderly women subjects with the combination of low appendicular skeletal muscle mass and high body fat mass is not yet known, and the association between other carotid artery parameters such as luminal diameter, flow velocity, and function with long-term exercise is not clear.

The aim of the present study was, therefore, to investigate the effect of combined exercise training on CIMT progression in SO elderly women. Identifying effective strategies for physical activity promotion to improve the risks of cardiovascular disease in elderly adults may reduce CVD, making this a key public health issue. We hypothesised that exercise training would reduce the progression of CIMT in elderly women with SO.

**Aim**

The purpose of this study was to determine the effects of a combined exercise program on carotid artery variables in elderly women with SO.

**Methods**

**Subjects**

This study was performed at the Buk-gu Sports Center and Institute of Taekwondo for Health and Culture, Dong-A University, between November 2015 and December 2016. The proposed study is a randomized single-blind controlled community-based trial with a parallel design and a 1:1 allocation ratio. We recruited women aged 65 years or older through recruitment notices in the community centers and health centers for the elderly in Busan City, South Korea. A total of 350 community-dwelling elderly women submitted applications to participate after reading recruitment notices. Inclusion criteria in this study were as follows: SO elderly women (≥ 65 years), Body mass index (BMI) ≥ 25.0 kg/m² and ASM/weight < 25.1 % were used as the criteria for SO in this study. Sixty elderly women were selected by the SO criteria after body composition measurement. Our subjects were nonsmokers in the past and present. Ten participants were excluded for the following reasons: history of CVD (n = 1) and medication for hypertension (n = 2), diabetes (n = 1), osteoporosis (n = 1), and arthritis (n = 1) and regular physical activity (n = 4, ≥ 20 min/day and ≥ 3 times/week). Finally, 50 subjects with SO were selected for this study. The 50 participants (aged 65 to 84 years) were divided into a control group (n = 25) and a supervised exercise program group (n = 25) by randomization (Fig. 1 and Table 1).

In accordance with the ethical standards of the Declaration of Helsinki, informed consent was obtained from all participants after they were provided with a detailed description of the experiment. Ethical approval was obtained from the ethics review board of Dong-A University.

**Body Composition and Blood Pressure**

After measuring the subjects’ height and body mass, their BFM percentage and ASM were measured by bioelectrical impedance analysis (BIA) using a body composition analyzer (Inbody 720, Biospace, Seoul, South Korea). The waist circumference (WC) was measured as the smallest horizontal girth between the costal margin and the iliac crests in the standing position. BMI (kg/m²) and ASM/weight were calculated with their respective formulas. Systolic blood pressure (SBP) and diastolic blood pressure (DBP) were measured using a mercury sphygmomanometer (HICO, Tokyo, Japan) after the subjects had rested for 10 minutes.

**Physical Function and Physical Activity**

Physical function was measured using muscle strength (MS), flexibility, walking speed, and aerobic endurance. MS was measured by a hand grip strength dynamometer (TKK-5401, Japan) and the 30-second chair stand-up test. Flexibility was measured with the sit-and-reach test. Maximum walking speed (MWS) was defined as the ratio between distance and time when walking 10 m. Aerobic endurance was measured with the 2-min step test. Physical activity was measured using the Korean short version of the international physical activity questionnaire (IPAQ). The IPAQ is available at https://sites.google.com/site/theipaq/.
Combined Exercise Program

The 24-week combined exercise program intervention consisted of sessions lasting 50–80 min, 5 days per week, of combined aerobic and resistance exercises under the supervision of an exercise specialist. The aerobic and resistance exercises were modified considering the characteristics of our subjects from physical activities recommended for the community-dwelling elderly23). Resistance exercises were performed with elastic band exercises (Thera-Band, Ohio, USA) for 12 items (elbow flexion, wrist flexion, shoulder flexion, lateral raise, front raise, chest press, reverse flies, side band, dead lift, squat, leg press, ankle plantar flexion), 8–15 repetitions per set (in weeks 1–12, 8–11 repetitions per set; in weeks 13–24, 12–15 repetitions per set), 2–3 sets (1 min rest between sets), 20–30 min per session for 3 days per week. Aerobic exercise involved various walking activities (sideways, backward, and forward walking and slow and fast indoor walking) for 30–50 min per session, 5 days per week, with a rating of perceived exertion (RPE) in the 13–17 range (in weeks 1–12, 13–15 RPE; in weeks 13–24, ≥15 RPE). Warm-up and cool-down was performed for 10 minutes before and after the exercise program, respectively. The control group was asked to maintain their usual physical activities during the 24 weeks, and we conducted health and family life education twice during the intervention period.

Blood Analysis

At baseline and after 24 weeks, venous blood samples were collected from both groups from an antecubital vein after an overnight fast (10–12 h). The concentrations of serum total cholesterol (TC), triglyceride (TG), high-density lipoprotein cholesterol (HDL-C), and low-density lipoprotein cholesterol (LDL-C) were analyzed using an automatic chemical analyzer (Hitachi-7600-110/7170 analyzer, Tokyo, Japan). High-sensitivity C-reactive protein (hs-CRP) level was measured by an automatic biochemical analyzer (Hitachi 747, Tokyo, Japan).

Carotid Artery Examination

CIMT, systolic and diastolic carotid artery luminal diameter (CLD), peak systolic flow velocity (PSV), and end diastolic flow velocity (EDV) were measured by B-mode ultrasound with a 10 MHz probe (LOGIQ 3, GE Healthcare, Wauwatosa, WI, USA). During measurement of the carotid arteries, the subjects lay on their backs in a dark room, heads turned 45 degrees, and the carotid arteries were fully exposed after relaxing for a minimum of 10 minutes; the left carotid artery was then measured by ultrasound. CIMT was measured on the far wall of the distal common carotid artery 1 cm proximal to the carotid bulb. CIMT was defined as the distance from the luminal-intima interface to the medial-adventitial interface. Systolic and diastolic CLD were measured at exactly the same location as CIMT measurement by B-mode ultrasound. CLD was defined as the distance from the intima-lumen interface of the near wall to the lumen-intima interface of the far wall. PSV and EDV were measured by continuous-wave Doppler examination in the carotid artery 1-3 cm proximal to the bifurcation9, 24). The sampling gate was placed at the center of the arterial axis, and FVs were recorded only after the signals stabilized. PSV and EDV were measured in the carotid artery with the insonation angle adjusted between 45° and 60° to the course of the vessel9, 20). Wall shear rate (WSR) was calculated using the Poi-
suillan parabolic model of velocity distribution across the arterial lumen on the basis of the assumption of laminar blood flow, according to the following formula: \( WSR_{\text{peak}} = 4 \times \frac{\text{PSV}}{\text{CLD}_{\text{systolic}}} \).

The test-retest coefficient of variation of CIMT, CLD, and PSV were 0.8%, 0.7%, and 0.7%, respectively. The intraclass correlation coefficient for repeated measures of the CIMT, CLD, and PSV were 0.7%, and echocardiographic measurements ranged from 0.6% to 0.9% in our laboratory.

### Statistical Analysis

Statistical Package for the Social Sciences (SPSS) ver. 17.0 (SPSS Inc., Chicago, IL, USA) was used for statistical analysis, and the measurement results are presented as averages and deviation. Student’s \( t \)-test was used to assess differences in baseline variables. Repeated two-way analysis of variance (ANOVA) was used to compare parameters before and after the 24 weeks of the study. If significant interactions were present, a paired sample \( t \)-test with Bonferroni correction was performed to identify differences of within-group factors. Statistical significance was set at \( p < 0.05 \).

### Results

Random grouping did not result in any significant differences in any variable between the groups at baseline (Table 1). All participants completed the intervention for 24 weeks. The mean combined exercise training attendance was 92% (86 to 100%).

Table 2 presents changes in body composition, blood pressure, physical function variables, and blood biochemical markers. In the ANOVA analysis results, BFM percentage \( (p < 0.001) \), WC \( (p = 0.002) \), left and right MS \( (p < 0.001) \), chair stand-up \( (p = 0.001) \), sit-and-reach \( (p = 0.003) \), MWS \( (p = 0.001) \), 2-min step \( (p < 0.001) \), SBP \( (p = 0.008) \), TC \( (p = 0.010) \), and LDL-cholesterol \( (p = 0.005) \) had significant interactions between group \( \times \) time. ASM, DBP, HDL-C, and hs-CRP had no interactions between group \( \times \) time. In the paired \( t \)-test results of the exercise group, BFM percentage \( (-2.0\%) \), WC \( (-0.7\,\text{cm}) \), SBP \( (-1.5\,\text{mmHg}) \), TC \( (-3.6\,\text{mg/dl}) \), and LDL-cholesterol \( (-3.7\,\text{mg/dl}) \) were significantly decreased, whereas left and right MS \( (2.5\,\text{kg}, 3.2\,\text{kg}) \), chair stand-up \( (1.5\,\text{repeated}) \), sit-and-reach \( (1.8\,\text{cm}) \), MWS \( (0.15\,\text{m/sec}) \), and 2-min walking \( (6.0\,\text{repeated}) \) were significantly increased. In the control group, left and right MS \( (-0.6\,\text{kg} and -0.5\,\text{kg}) \), chair stand-up \( (-1.0\,\text{repeated}) \), MWS \( (-0.03\,\text{m/sec}) \), and 2-min step \( (-1.9\,\text{repeated}) \) were significantly decreased.

Table 3 presents changes of carotid artery parameters. In the ANOVA analysis results, CIMT \( (p = 0.013) \), PSV \( (p = 0.007) \), EDV \( (p = 0.006) \), and WSR \( (p < 0.001) \) showed significant group \( \times \) time interactions. CLDs (both systolic and diastolic) had no group \( \times \) time interactions. In the paired \( t \)-test results of the exercise group, CIMT \( (-0.01\,\text{mm}) \) was significantly decreased, whereas PSV \( (1.4\,\text{cm/sec}) \), EDV \( (0.3\,\text{cm/sec}) \), and WSR \( (10.5\,\text{s}^{-1}) \) were significantly increased. However, CLDs (both systolic and diastolic) in the exercise group showed no change nor did any of the carotid artery parameters in the control group.

### Discussion

This study examined the effects of a 24-week combined exercise program on carotid artery parameters in elderly women with SO. The main finding of this study is that the 24-week combined exercise program reduced CIMT in elderly women with SO. Additionally, the second finding of our study is that long-term regular exercise increased PSV and WSR in elderly women.
SO is more closely associated with traditional CVD risk factors when compared with either obesity or sarcopenia in the elderly\(^1\). Further, SO is more closely associated with increasing CIMT when compared with either obesity or sarcopenia\(^8\). Even though previous evidence for exercise and CIMT suggests that exercise training does not affect CIMT in healthy young men and women\(^17-19\), two studies of association between exercise training and CIMT demonstrated that aerobic exercise training for 12 or 24 weeks reduced CIMT in overweight and obese adults (-7.8%) and formerly preclamptic women (-8.6%)\(^25, 27\). Moreover, one intervention study reported that aerobic exercise for 24 weeks is effective in decreasing the artery wall thickness in a large artery in overweight and obese elderly adults\(^28\). In our study, CIMT was decreased (-1.5%) through combined exercise for 24 weeks; however, the effect of exercise training on CIMT was smaller than it was in two previous intervention studies\(^26, 27\).

A recent intervention study by Byrkjeland et al. demonstrated that combined exercise training is effective in decreasing CIMT (-0.034 mm) in high-risk individuals with the combination of type 2 diabetes and coronary artery disease (CAD)\(^29\). Our study subjects were elderly women with sarcopenic obesity. Obesity and sarcopenia are proven risk factors for CVD and disability in the elderly\(^30, 31\). Our study

### Table 2. Changes in the body composition, physical function, blood pressure, serum lipid and high-sensitivity C-related protein

<table>
<thead>
<tr>
<th></th>
<th>Control (n = 25)</th>
<th>Exercise (n = 25)</th>
<th>F-values</th>
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<tbody>
<tr>
<td></td>
<td>Baseline</td>
<td>24 week</td>
<td></td>
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<tr>
<td>Percentage body fat (%)</td>
<td>40.4 ± 3.6</td>
<td>40.8 ± 4.2</td>
<td></td>
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<tr>
<td>Waist circumference (cm)</td>
<td>90.2 ± 4.2</td>
<td>90.4 ± 4.1</td>
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<tr>
<td>ASM (kg)</td>
<td>15.0 ± 2.0</td>
<td>14.9 ± 1.6</td>
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<td>Left grip strength (kg)</td>
<td>22.5 ± 3.1</td>
<td>21.9 ± 2.7*</td>
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<tr>
<td>Right grip strength (kg)</td>
<td>23.3 ± 2.9</td>
<td>22.8 ± 2.8*</td>
<td></td>
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<tr>
<td>CSU (repetitions/30 sec)</td>
<td>16.6 ± 2.9</td>
<td>15.6 ± 2.4</td>
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<tr>
<td>Sit and reach (cm)</td>
<td>12.9 ± 4.7</td>
<td>12.4 ± 3.9</td>
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<tr>
<td>MWS (m/sec)</td>
<td>1.47 ± 0.21</td>
<td>1.43 ± 0.21**</td>
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<tr>
<td>Two-minute step (repetitions)</td>
<td>98.5 ± 9.0</td>
<td>96.6 ± 10.2*</td>
<td></td>
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<td>SBP (mmHg)</td>
<td>128.6 ± 7.3</td>
<td>128.9 ± 6.8</td>
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<tr>
<td>DBP (mmHg)</td>
<td>78.2 ± 6.1</td>
<td>78.6 ± 4.6</td>
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<td>Total cholesterol (mg/dl)</td>
<td>187.7 ± 10.8</td>
<td>187.6 ± 9.5</td>
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<tr>
<td>Triglyceride (mg/dl)</td>
<td>116.9 ± 10.7</td>
<td>116.2 ± 10.4</td>
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<tr>
<td>LDL- cholesterol (mg/dl)</td>
<td>100.7 ± 14.9</td>
<td>102.4 ± 11.8</td>
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<tr>
<td>HDL- cholesterol (mg/dl)</td>
<td>49.0 ± 6.9</td>
<td>49.1 ± 5.5</td>
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<tr>
<td>High-sensitivity CRP (mg/L)</td>
<td>0.12 ± 0.13</td>
<td>0.12 ± 0.13</td>
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</table>

Data are show as mean ± SD. ASM, appendicular skeletal muscle mass; CSU, chair stand up; MWS, maximum walking speed; SBP, systolic blood pressure; DBP, diastolic blood pressure; LDL, low density lipoprotein; HDL, high density lipoprotein; CRP, C-reactive protein; \(^* p<0.05, \^* p<0.01, \^\* p<0.001, \text{change within the groups}; \^ p<0.05, \^\* p<0.01, \^\*\* p<0.001, group \times time interaction

### Table 3. Changes in the carotid artery intima-media thickness, luminal diameter, flow velocity and wall shear rate

<table>
<thead>
<tr>
<th></th>
<th>Control (n = 25)</th>
<th>Exercise (n = 25)</th>
<th>F-values</th>
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</thead>
<tbody>
<tr>
<td></td>
<td>Baseline</td>
<td>24 week</td>
<td></td>
</tr>
<tr>
<td>CIMT (mm)</td>
<td>0.67 ± 0.07</td>
<td>0.67 ± 0.07</td>
<td></td>
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<tr>
<td>Diastolic CLD (cm)</td>
<td>0.63 ± 0.09</td>
<td>0.63 ± 0.09</td>
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<tr>
<td>Systolic CLD (cm)</td>
<td>0.58 ± 0.08</td>
<td>0.58 ± 0.07</td>
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<tr>
<td>PSV (cm/sec)</td>
<td>64.2 ± 10.2</td>
<td>64.3 ± 9.5</td>
<td></td>
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<tr>
<td>EDV (cm/sec)</td>
<td>19.8 ± 4.9</td>
<td>19.8 ± 4.9</td>
<td></td>
</tr>
<tr>
<td>Wall shear rate (s(^{-1}))</td>
<td>454.7 ± 110.2</td>
<td>452.7 ± 103.3</td>
<td></td>
</tr>
</tbody>
</table>

Data are show as mean ± SD. CIMT, carotid artery intima-media thickness; CLD, carotid artery luminal diameter; PSV, peak systolic flow velocity; EDV, end diastolic flow velocity; \(^* p<0.05, \^* p<0.01, \^\* p<0.001, \text{change within the groups}; \^ p<0.05, \^\* p<0.01, \^\*\* p<0.001, group \times time interaction
results imply that 6 months of combined exercise is effective in reducing CIMT in elderly women with a combination of low skeletal muscle mass and high body fat mass. Nevertheless, we believe that the effects of exercise on CIMT will appear in limited subjects. Recent reviews on exercise training and atherosclerosis have reported that the reduction in CIMT as a result of exercise training is associated with increased cyclic blood pressure, shear stress, and antioxidant defense and decreased inflammatory processes, vascular tone, and sympathetic nervous system activity. Another review reported that increased physical activity suppresses the overall CVD risk and hence curtails the progression of carotid atherosclerosis. The rationale for the association between exercise and CIMT presented in the two reviews is considered to be a potential mechanism for the reduction of CIMT based on combined exercise in this study.

A cross-sectional study and a long-term follow-up study reported that a large CLD and low FV are closely associated with CVD and stroke in healthy subjects and in patients. Another cross-sectional study reported that CLD in elderly people with high cardiorespiratory function, as evaluated by oxygen uptake, is narrower than it is in those with low cardiorespiratory function. However, the evidence for the association of CLD and FV with exercise training is very limited. A few interventional studies have reported that short-term exercise training does not affect the CLD and FV in healthy young men. In our study, after combined exercise for 24 weeks, CLDs showed no change; however, PSV (2.1%), EDV (1.5%), and WSR (2.3%) were significantly increased in the exercise group.

Low FV of the carotid artery is associated with BFM and other CVD risk factors. Low FV and WSR are also associated with accelerated atherosclerosis progression. Therefore, this study found that regular exercise increases carotid artery FV and WSR in elderly women with SO. The mechanism for changes in the FV in large arteries is not clear. However, several previous studies reported that resting cardiac output is related to middle cerebral artery FV, and regular exercise improves cardiac function readings such as cardiac output, stroke volume, and ejection fraction. Based on these facts, we think that the improvement of cardiac function such as cardiac output and stroke volume by exercise training contributed to the increase in carotid artery FV. Due to our hypothesis on the mechanism of FV change by exercise training, we have limitations in not investigating cardiac function. In the present study, PSV increased, but the systolic CLD did not change. The change in WSR is determined by changes in PSV and systolic CLD. Therefore, the change in WSR is thought to be due to the increase in PSV. Additionally, the decreasing tendency of systolic CLD may have contributed a little to the increase in WSR. The results of this study suggested that increases in carotid artery FV and WSR by exercise training are considered to have additional benefit for CVD prevention in elderly women subjects with SO. However, more evidence is required to prove a clear association between the carotid FV and regular exercise.

Combined exercise training is effective in improving body composition, physical function, blood pressure, and blood biochemical markers. Resistance exercise in previous studies used a special exercise machine, requiring a special place and providing similar effects on body composition and physical function. Therefore, resistance exercise was performed with an elastic band, and aerobic exercise was performed through walking. Our combined exercise improved the body composition and physical function in elderly women.

Several limitations were inherent in this study. ASM was evaluated by BIA in this study. ASM evaluation by the BIA method has a higher correlation with dual-energy X-ray (DXA), and this method is simple and inexpensive when compared with the DXA method in ASM evaluation. Therefore, BIA was used to evaluate body composition. However, DXA is more widely used in evaluating ASM. Before interpreting the results, it should be noted that there are certain facts to be considered in this research. The time of the daily physical activities during intervention was not measured to mention that the diet of either group was not controlled. Moreover, the sample size was small, which is the significant determinant of the research; each factor could have altered the result of the study. Hence, in order to understand the impact of exercise on carotid artery parameters, a bigger sample size and a longer period of trial are needed.

**Conclusions**

In conclusion, the 24-week combined exercise program effectively decreased CIMT and increased carotid artery FV as well as WSR in elderly women with SO. Therefore, regular combined exercise is thought to contribute to a lower CVD risk in elderly women with SO.

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Disclosure

We have no conflict of interest or financial disclosures to declare in conjunction with the publication of this work.

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