Original/Deporte y ejercicio

Positive effects of resistance training on inflammatory parameters in men with metabolic syndrome risk factors

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Summary

Background: evidences have shown a strongly association between metabolic syndrome (MS), cardiovascular diseases and chronic low-grade inflammation, being this last, related with the occurrence of sarcopenia and atherosclerosis. Despite several benefits, the effects of resistance training (RT) on inflammatory profile are controversial. Thereby, this study aims to investigate the effects of a RT on the inflammatory profile of men with MS risk factors.

Methods: fifteen sedentary men (57.53 ± 7.07 years old) with 2 or more MS components underwent a RT for 14 weeks (3 times per week), with intensity ranging between 40 and 70% of one repetition maximum. The dual-energy X-ray absorptiometry was used to body composition assessment and serum was collected to evaluate biochemical and inflammatory parameters before and after the RT.

Results: despite the absence of changes in body weight, total muscular content and biochemical parameters, the individuals demonstrated a reduction on body fat content (p < 0.05). Furthermore, the RT resulted in lower circulating levels of tumor necrosis factor alpha and interleukin-6 (p < 0.05), in higher levels of interleukin-10 (p < 0.05) and in the stabilization of interleukin-1 beta and interferon-gamma concentrations. It was concluded that a moderate RT benefits inflammatory profile, contributing to a lower risk of cardiovascular diseases.

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Key words: Health. Cardiovascular diseases. Exercise. Cytokines. Adiposity.

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EFECTOS POSITIVOS DEL ENTRENAMIENTO DE RESISTENCIA EN LOS PARÁMETROS INFLAMATORIOS DE HOMBRES CON FACTORES DE RIESGO DE SÍNDROME METABÓLICO

Resumen

Introducción: la evidencia muestra la relación entre síndrome metabólico (SM), enfermedades cardiovasculares e inflamación crónica de bajo grado, que está relacionada con la aparición de sarcopenia y aterosclerosis. A pesar de muchos beneficios, los efectos del entrenamiento de resistencia (ER) sobre la inflamación son controvertidos. Este estudio tiene como objetivo investigar los efectos de 14 semanas de ER en el perfil inflamatorio de hombres con factores de riesgo para SM.

Métodos: quince varones sedentarios (57,53±7,07 años) con 2 o más componentes del SM fueron sometidos a un ER moderado durante 14 semanas (tres veces a la semana), con una intensidad que oscila entre 40 y 70% de una repetición máxima. La absorciometría dual de rayos X se utilizó para la evaluación de la composición corporal y el suero se recogió para evaluar los parámetros bioquímicos e inflamatorios antes y después de la ER.

Resultados: a pesar del mantenimiento de la masa corporal, la masa muscular total y los parámetros bioquímicos, hubo una reducción en la grasa corporal (p<0,05). Además, el ER disminuyó los niveles de factor de necrosis tumoral alfa e interleucina-6 circulante (p<0,05), aumentó la concentración de interleucina-10 (p<0,05) y mantuvo la interleucina-1 y el interferón-gama. Se concluyó que 14 semanas de ER moderado provocan beneficios sobre el perfil inflamatorio, contribuyendo a la reducción del riesgo cardiovascular.

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Functional Assessments

All the tests described below were performed by the same researcher and at the same time of day, before and after the 14th week of RT. To estimate the largest load that an individual can move in a single maximal effort and thus prescribe the training load, a submaximal test was used to estimate the 1RM in the bench press, rower machine, leg press, knee flexion exercises. Cardiorespiratory fitness was assessed by Bruce’s modified protocol in a treadmill. The levels of systolic blood pressure (SBP) and diastolic blood pressure (DBP) were measured after the subjects sat quietly for 5 min with a digital sphygmomanometer (Omron, Kyoto, Japan). Furthermore, the flexibility of lumbar and hamstring muscles were assessed by the sit-and-reach test and the longest distance reached on the measuring board was registered after three attempts.

Anthropometric Measurements

The subjects were weighted in a scale (Plenna, São Paulo, Brazil) and height was evaluated with a stadiometer (Cardiomed, Curitiba, Brazil). The abdominal circumference was measured with a spring-loaded metal tape (Cardiomed, Curitiba, Brazil). Body composition was determined using dual-energy X-ray absorptiometry (DXA) with a densitometer machine (Hologic QDR Discovery, Waltham, USA). Briefly, after 12 h fasting and 24 h without exercises and wearing only a light coat, subjects lay supine on the DXA table with their arms adequately separated from their trunk and were instructed to remain still throughout the scanning procedure.

Biochemical Assays

Blood samples were collected from a vein of the antecubital region after 12 h of fasting and 72 h without physical exercise practice. Samples were collected into serum separator tubes (BD Diagnostics, Plymouth, United Kingdom) and centrifuged at 1500 g for 15 min. Then, serum was frozen at -80ºC until analysis. The triglycerides, total cholesterol, HDL and glucose levels were determined spectrophotometrically with commercially available assay kits (Labtest, Lagoa Santa, Brazil). The concentration of low-density cholesterol (LDL) was estimated. The serum levels of cytokines IL-1β, IL-6, IL-10, TNF-α and interferon-gamma (IFN-γ) were determined by enzyme-linked immunosorbent assay (ELISA) using commercial kits for human (eBIOSCIENCE, San Diego, USA), according to manufacturer’s instructions. Lipid and glucose parameters were analyzed with samples in duplicate, while inflammatory parameters were performed with samples in triplicate.

Nutritional Data

To minimize a possible bias, subjects were encouraged to not alter their habitual dietary intake during the intervention and completed a 3-day diet record in the beginning and in the end of the RT. The 3-day diet record included Sunday, Monday and Tuesday analysis. To determine total caloric intake and the amount of macronutrients ingested, it was utilized a specific software (Dietwin, São Paulo, Brazil).

Statistical Analysis

The Shapiro-Wilk test was carried out to verify data normality. Afterwards, Student’s t test or Wilcoxon Rank Test were used to determine significant differences between the pre and post-training means. Statistical Package for Social Sciences (SPSS 14.0, Chicago, USA) was used and statistical significance was set at p < 0.05. Data were expressed as mean ± standard deviation of the mean (SD).

Results

Of the 18 volunteers who started participating in the study, 15 individuals (57.53 ± 7.07 years old) concluded the RT and were considered in the statistical analysis (three individuals were excluded due the absence in more than 25% of RT sessions). Besides, the sample comprised two smokers, six former smokers and seven nonsmokers. Furthermore, 40% of men took antihypertensive agents, 33.3% took lipid-lowering agents and 20% took oral hypoglycemic agents. Among the fifteen subjects, the adherence to RT was of 82.34%. In Table I, it is demonstrated that TR did not result in significant changes in most anthropometric, functional and biochemical characteristics. However, total body fat content decreased with the RT (p = 0.041).

Table II exhibits the results of the submaximal strength test before and after the RT. Increases in the load lifted/moved in the bench press (p < 0.001), leg press (p < 0.001), rower machine (p < 0.001) and knee flexion (p = 0.003) exercises were observed.

No significant differences were reported in the total caloric intake and the amount of macronutrients ingested of carbohydrates, proteins and lipids, demonstrating the maintenance of habitual intake during the intervention (Table III).

Figure 1 shows the impact of RT on serum levels of cytokines. It was observed that RT decreased TNF-α (174.2 ± 56.59 vs. 140 ± 23.25 pg/mL; p = 0.011) (B), IL-6 (155 ± 50.18 vs. 124 ± 26.73 pg/mL; p = 0.019) (D), and IL-10 levels (86.53 ± 34.93 vs. 108.53 ± 24.76 pg/mL; p = 0.037) (E). However, the IL-1β (p = 0.074) (A) and INF-γ levels (p = 0.084) (C) did not change significantly after the intervention.
This study aimed to investigate the effects of a supervised RT on inflammatory profile in men with metabolic syndrome risk factors. The main findings were that RT provoked improvements in inflammatory and anthropometric (body fat content) parameters, independently of significant changes in body mass and in the factors of MS classification. Furthermore, RT promoted increases in exercise load during 1RM test, indicating a functional adaptation to the stimulus generated from RT sessions.

Evidences have demonstrated that the age-related skeletal muscle catabolism and sedentary lifestyle contribute to the high prevalence of metabolic disorders. In this sense, a recent review and meta-analysis demonstrated that RT, as a single intervention, is not able to change total cholesterol, triglycerides, HDL, LDL and diastolic blood pressure, as the results obtained in this study. Furthermore, beneficial alterations in inflammatory biomarkers associated with the physical training are not dependent of weight loss, conflicting with some previous studies.

Although the regular practice of resistance exercise provides several health benefits, the effects of RT on cytokines parameters are inaccurate. In the present study, 14 weeks of moderate RT resulted in lower levels of serum TNF-α and IL-6 and higher levels of IL-10 in men with MS, while IL-1β and IFN-γ levels was not altered. However, other investigations with RT protocols in individuals with considerable cardiovascular risk did not find changes in circulating inflammatory markers after intervention. Possible reasons for conflicting results may include age of the participants, basal levels of cytokines, influence of the last exercise session, besides frequency, duration and intensity of training.

Studies demonstrate that positive effects of RT on inflammatory profile vary according to the biomarker considered.

### Table I

<table>
<thead>
<tr>
<th>Parameters</th>
<th>Before</th>
<th>After</th>
</tr>
</thead>
<tbody>
<tr>
<td>Body Mass (kg)</td>
<td>89.24 ± 15.5</td>
<td>88.82 ± 14.77</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>29.82 ± 5</td>
<td>29.68 ± 4.81</td>
</tr>
<tr>
<td>Abdominal Circumference (cm)</td>
<td>106.47 ± 11.34</td>
<td>105.28 ± 10.66</td>
</tr>
<tr>
<td>Total Body Fat Content (kg)</td>
<td>29.78 ± 9.19</td>
<td>28.57 ± 8.71</td>
</tr>
<tr>
<td>Total Muscle Mass (kg)</td>
<td>54.91 ± 5.56</td>
<td>55.29 ± 5.99</td>
</tr>
<tr>
<td>Systolic Blood Pressure (mmHg)</td>
<td>122.2 ± 20.26</td>
<td>118.33 ± 16.86</td>
</tr>
<tr>
<td>Diastolic Blood Pressure (mmHg)</td>
<td>73.66 ± 9.65</td>
<td>69.80 ± 9.83</td>
</tr>
<tr>
<td>VO₂max (mL/kg⁻¹·min⁻¹)</td>
<td>32.29 ± 10.32</td>
<td>33.47 ± 11.34</td>
</tr>
<tr>
<td>Glucose (mg/dL)</td>
<td>96.03 ± 11.85</td>
<td>105.87 ± 23.36</td>
</tr>
<tr>
<td>Total Cholesterol (mg/dL)</td>
<td>156.38 ± 41.09</td>
<td>166.37 ± 45.68</td>
</tr>
<tr>
<td>Triglycerides (mg/dL)</td>
<td>162.46 ± 71.23</td>
<td>159.46 ± 90.37</td>
</tr>
<tr>
<td>HDL (mg/dL)</td>
<td>42.9 ± 14.91</td>
<td>46.78 ± 17.98</td>
</tr>
<tr>
<td>LDL (mg/dL)</td>
<td>80.99 ± 31.52</td>
<td>80.19 ± 49.09</td>
</tr>
</tbody>
</table>

Values expressed as mean ± SD. BMI: body mass index. VO₂max: maximal oxygen uptake. HDL: high-density cholesterol. LDL: low-density cholesterol. *p < 0.05 after vs. before resistance training.

### Table II

<table>
<thead>
<tr>
<th>Exercises</th>
<th>Before</th>
<th>After</th>
</tr>
</thead>
<tbody>
<tr>
<td>Bench Press (kg)</td>
<td>43.98 ± 11.40</td>
<td>61.36 ± 15.90**</td>
</tr>
<tr>
<td>Leg Press (kg)</td>
<td>98.28 ± 22.67</td>
<td>122.20 ± 20.09**</td>
</tr>
<tr>
<td>Rower machine (kg)</td>
<td>44.76 ± 13.80</td>
<td>57.18 ± 9.80**</td>
</tr>
<tr>
<td>Knee Flexion (kg)</td>
<td>27.22 ± 7.16</td>
<td>41.6 ± 13.65*</td>
</tr>
</tbody>
</table>

**p < 0.01 and *p < 0.01 after vs. before the resistance training.

### Table III

<table>
<thead>
<tr>
<th>Variables</th>
<th>Before</th>
<th>After</th>
</tr>
</thead>
<tbody>
<tr>
<td>Total Caloric Intake (kcal)</td>
<td>2171.14 ± 461.17</td>
<td>2137.93 ± 494.78</td>
</tr>
<tr>
<td>Carbohydrates (g)</td>
<td>242.97 ± 51.54</td>
<td>233.89 ± 65.19</td>
</tr>
<tr>
<td>Proteins (g)</td>
<td>97.31 ± 20.86</td>
<td>106.16 ± 22.05</td>
</tr>
<tr>
<td>Lipids (g)</td>
<td>94.42 ± 25.50</td>
<td>88.50 ± 24.85</td>
</tr>
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</table>

Values expressed as mean ± SD.

### Discussion

This study aimed to investigate the effects of a supervised RT on inflammatory profile in men with metabolic syndrome risk factors. The main findings are that RT provoked improvements in inflammatory and anthropometric (body fat content) parameters, independently of significant changes in body mass and in the factors of MS classification. Furthermore, RT promoted increases in exercise load during 1RM test, indicating a functional adaptation to the stimulus generated from RT sessions.

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Studies demonstrate that positive effects of RT on inflammatory profile vary according to the biomarker considered.
kers. For instance, four months of RT with post-menopausal women decreased IL-6 levels, despite the maintenance of TNF-α and interleukin-15 concentrations\cite{15}. Similarly, 12 weeks of RT lowered TNF-α, leptin and C reactive protein (CRP) levels, although the maintenance of other plasma and subcutaneous adipose tissue inflammatory markers levels in postmenopausal women\cite{14}. However, 32 weeks of resistance exercise decreased IL-6 levels in elderly men, without modifying IFN-γ, TNF-α and CRP concentrations\cite{29}. In this regard, protocols of RT with lower week frequency and amount of exercises induced decreasing of CRP, IFN-γ, serum amyloid A and heat shock protein 70, respectively, despite the unchanged levels of TNF-α and IL-6\cite{16,30}.

Considering that adipose tissue is an endocrine organ\cite{4}, a reduction in the adipose tissue content may influence the production and releasing of proinflammatory markers, as confirmed in the present study. It is highlighted that the found body fat content reduction derives from the RT, since caloric intake and amount of macronutrients did not change along the intervention. The IL-6 releasing from adipose tissue and immune cells is linked with several disorders, characterizing the chronic low-grade inflammation status. In contrast, investigations have demonstrated that IL-6 released by skeletal muscle during the physical exercise stimulates the lipolysis, the IL-10 production and inhibits the TNF-α production\cite{4}. In this sense, in vivo and in vitro studies have demonstrated that IL-6 release from myocytes is an essential regulator of skeletal muscle hypertrophy mediated by satellite-cells\cite{31}. The physical training also reduces skeletal muscle mRNA toll-like receptor 4 (TLR4) and TNF-α, possibly influencing circulating cytokines levels.

Other several mechanisms have been proposed to explain the positive effects of RT on inflammatory parameters in previously sedentary individuals. It has been suggested that anti-inflammatory effects of RT are mediated by the modulation of cytokines production not only in the adipose tissue, but also in the skeletal muscle and peripheral blood mononuclear cells\cite{15}. Another hypothesis is that RT results in reduced expression of TLR4 in monocytes, as previously demonstrated\cite{33}. Indeed, TLR4 is known to activate the nuclear factor kappa B (NFκB), inducing the expres-

![Fig. 1.—Effects of resistance training on circulating levels of interleukin-1 beta (IL-1β) (A), tumor necrosis factor alpha (TNF-α) (B), interferon-gamma (INF-γ) (C), interleukin-6 (IL-6) (D), e interleukin-10 (IL-10). * p < 0.05 after vs. before the resistance training.](image-url)
sion of proinflammatory cytokines such TNF-α and IL-6\(^4\). Furthermore, investigations have demonstrated that physiological adjustments to muscle contraction involves functional changes in macrophages, reducing the production of TNF-α and IL-1β due anti-inflammatory cytokines, such as IL-10\(^3\),\(^5\). In this regard, it is known that higher levels of proinflammatory cytokines stimulate the migration of monocytes to the arterial layer and its conversion in macrophages, originating atherosclerosis lesions\(^9\).

In conclusion, these results suggest that 14 weeks of moderate resistance training induces positive and favorable effects on inflammatory profile in men with metabolic syndrome risk factors, contributing to the decrease of cardiovascular risk. The lack of control group may be considered a limitation of this study.

**Disclosure of interest**

The authors declare that they have no conflicts of interest concerning this article.

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