Is dynamometry able to infer the risk of muscle mass loss in patients with COPD?

Introduction: Sarcopenia is characterized by a progressive and generalized decrease of strength and muscle mass. Muscle mass loss is prevalent in patients with chronic obstructive pulmonary disease (COPD) as a result of both the disease and aging. Some methods have been proposed to assess body composition (and therefore identify muscle mass loss) in this population. Despite the high accuracy of some methods, they require sophisticated and costly equipment.

Aim: The purpose of this study was to infer the occurrence of muscle mass loss measured by a sophisticated method (dual energy X-ray absorptiometry [DEXA]) using a more simple and affordable equipment (dynamometer).

Methods: Fifty-seven stable subjects with COPD were evaluated for anthropometric characteristics, lung function, functional exercise capacity, body composition, and peripheral muscle strength. A binary logistic regression model verified whether knee-extension strength (measured by dynamometry) could infer muscle mass loss (from DEXA).

Results: Patients with decreased knee-extension strength were 5.93 times more likely to have muscle mass loss, regardless of sex, disease stage, and functional exercise capacity ($P=0.045$).

Conclusion: Knee-extension dynamometry was able to infer muscle mass loss in patients with COPD.

Keywords: COPD, sarcopenia, peripheral muscle strength

Chronic obstructive pulmonary disease (COPD) is characterized by a variety of pathophysiological conditions and clinical manifestations that contribute to the heterogeneity of patient profiles. This disease is a major cause of chronic morbidity and the fourth leading cause of mortality worldwide. It is estimated that, in 2020, it will be the third highest cause of death worldwide. Among the systemic manifestations observed in COPD are the progressive decrease of skeletal muscle mass and the presence of several bioenergetic abnormalities, mainly expressed by low body weight. The disease’s systemic effects can greatly increase the clinical symptoms, such as limitation of exercise capacity and peripheral muscle weakness and can have a negative impact on quality of life. Weight loss and low body mass index (BMI) have been suggested to be associated with worse health status and higher mortality rates in these patients.

The aging process initiates sarcopenia, a syndrome characterized by progressive decrease of strength and muscle mass. Muscle mass loss is prevalent in patients with COPD as a result of both the disease and aging. Pulmonary rehabilitation programs are aimed at maintenance or improvement of physical and nutritional condition of individuals with COPD. In this sense, the...
assessment of patients’ peripheral muscle strength and body composition is important to provide them an appropriate treatment and follow-up their progress during pulmonary rehabilitation programs.\textsuperscript{14}

A few methods have been proposed to assess body composition in this patient population.\textsuperscript{15} Despite some of them being highly accurate, they require sophisticated and costly equipment, such as Dual Energy X-Ray Absorptiometry (DEXA), restricting its applicability in the clinical practice. Thus, from a clinical perspective, a method should be practical, portable, and inexpensive.\textsuperscript{16–18}

Given the negative impact on quality of life that peripheral muscle strength and muscle mass may have due to the presence of systemic alterations, there is a clear need for studies proving the benefits of simple and accessible assessment tools that can be widely used in clinical practice in patients with COPD.

The purpose of this study was to verify whether it was possible to infer the occurrence of muscle mass loss measured by a sophisticated method (DEXA) using a more simple and affordable equipment (dynamometer).

\section*{Methods}

\subsection*{Participants}

The study participants were patients with COPD referred to the Pulmonary Rehabilitation Center for Studies, of the Faculty of Sciences and Technology, UNESP, Presidente Prudente, São Paulo, Brazil. The criteria for participant selection were diagnosis of COPD according to the (Global initiative for chronic Obstructive Lung Disease) GOLD criteria\textsuperscript{19} and clinical stability with no exacerbations or changes in medication for at least 30 days. The exclusion criterion was the presence of disabling pathological conditions that could potentially influence physical activity performance, such as cerebrovascular diseases, rheumatism, and arthritis. A total of 57 patients were included in the study.

The study was approved by the institutional review board, the Ethics Committee of São Paulo State University/ Presidente Prudente, (Protocol number 95/2009), and all individuals signed an informed consent before participating in the study.

\subsection*{Outcome measures}

Patients included in the study were evaluated for their anthropometric characteristics, lung function, functional exercise capacity, body composition, and peripheral muscle strength.

Body weight was measured on a digital balance accurate to 0.1 kg (Welmy R/I 200; Welmy, São Paulo, Brazil). Height was determined using a stadiometer accurate to 1 cm (Sanny; American Medical Do Brasil Ltda, São Bernardo do Campo, Brazil). BMI was calculated as the ratio of body weight (kg) and height squared (cm\textsuperscript{2}) \((\text{BMI} = \text{body weight/height}^2)\).

Lung function was assessed by spirometry using a portable spirometer Spirobank, version 3.6: (MIR, Rome, Italy) and performed according to the guidelines of the American Thoracic Society and European Respiratory Society.\textsuperscript{20} The reference values were specific for the Brazilian population.\textsuperscript{21}

The 6-minute walk test (6MWT) was used to assess functional exercise capacity according to the guidelines of the American Thoracic Society.\textsuperscript{22} Results obtained were referred and compared to normal values.\textsuperscript{23}

Peripheral muscle strength tests of knee flexion and extension, shoulder flexion and abduction, and elbow flexion were performed unilaterally (dominant limb) with a digital dynamometer (Force Gauge® brand, FG -100 kg, Sao Paulo, Brazil), and the results were expressed in Newton (N). The volunteers performed a maximal voluntary isometric contraction for 6 seconds followed by limb relaxation. The measurement was repeated thrice with intervals of 1 minute, and the highest value was recorded.

The lean mass was assessed by DEXA (Hologic model QDR machine dials 2000/Plus; Hologic, Waltham, MA, USA) Software version 5.56.\textsuperscript{24} The ratio of appendicular lean mass (upper limbs lean mass + lower limbs lean mass [kg]/height [m\textsuperscript{2}]) was calculated and used to classify subjects into high lean mass (HLM) or low lean mass (LLM). Men with values lower than 7.91 kg/m\textsuperscript{2} and women with values lower than 5.52 kg/m\textsuperscript{2} were considered sarcopenic and included in the LLM. The adoption of these cut-off points was based on two standard deviations below the mean of a youth group (n=30, 20 men and ten women) aged between 20 and 30 years old, according to the recommendations of Baumgartner et al.\textsuperscript{25} Therefore, values below 206.4 and 144.5 N for knee extension, 112 and 72.7 N for knee flexion, 62.9 and 39 N for shoulder flexion, 58.8 and 33.6 N for shoulder abduction, and 116 and 59.2 N for elbow flexion, for men and women, respectively were considered as LLM.

\subsection*{Data analysis}

Data were presented as mean and standard deviation. Data distribution was tested by the Shapiro–Wilks test. The mean values for each variable were compared between the groups (HLM and LLM) by the unpaired Student’s \(t\)-test.
The chi-square test was used to compare the proportion of patients above and below the cut-offs of muscle strength (see “Methods” section) in LLM. A binary logistic regression model was created to express the magnitude of the associations by odds ratio values (OR). In this model, the outcomes of muscle strength, sex, disease severity (GOLD), and functional exercise capacity were set as independent variables. Then, we verified the association between the variables in the model and each of the five tested muscle groups (knee extension/flexion, shoulder flexion/abduction, and elbow flexion). Statistical analysis was performed using SPSS software (SPSS Inc., Chicago, IL, USA) version 17.0, and significance level was set at 5%.

Results
Fifty-seven patients participated in the study, of them 20 were women (forced expiratory volume in the first second [FEV₁], 44±14.1 percentage of predicted values [%pred]) and 37 were men (FEV₁, 52.6±19.9 %pred). There was no significant difference in spirometric characteristics between men and women (P=0.091).

Patients’ anthropometric characteristics, their values of upper and lower limbs muscle strength, and the comparison between HLM and LLM are shown in Table 1. Twenty-five patients (19 men and six women) had sarcopenia and were included in the LLM group. Men with LLM had lower weight, BMI, appendicular lean mass, and lower knee extension strength than nonsarcopenic individuals (P<0.01 for all). In the LLM women group, there was lower appendicular lean mass than in HLM group (P=0.011).

The results from the association between muscle strength tests for each movement and LLM can be seen in Table 2. Only knee extension presented a significant association. Patients with decreased knee extension strength were 5.93 times more likely to have low lean mass, regardless of sex, disease stage, and functional exercise capacity (P=0.045).

Discussion
The present investigation showed that the use of a dynamometer to assess peripheral muscle strength, specifically the knee-extension strength, was able to infer muscle mass loss in patients with COPD.

Few studies have focused on the relationship between muscle strength and muscle mass in subjects with COPD. Villaca et al.²⁶ correlated dynamometry and the leg lean volume measured by DEXA. Hillman et al.²⁷ investigated the relationships of body composition and peripheral muscle strength (measured by dynamometry) with 6-minute walk distance. This is the first study that aimed at verifying the extent to which it was possible to infer muscle lean mass based on muscle strength.

We observed that knee-extension strength was highly associated with muscle mass depletion. Studies have demonstrated a relationship between peripheral muscle dysfunction and local tissue depletion in patients with COPD.¹⁵,²⁶–³⁰ Franssen et al.¹⁰ described different adaptations in muscles of lower and upper limbs after a training program. Furthermore, it is also described that individuals may have lower lean mass index with preserved muscle strength.²⁵ This may explain why we only found significant association with lower limb force and not with any of the tested upper limb groups.

Hillman et al.²⁷ observed a correlation between the quadriceps strength measured with a dynamometer and the lower limb muscle mass measured by DEXA. In their study, it was shown that the measure of muscle mass by DEXA was not able to add clinical information on muscle function from that observed by the simple measure from a

Table 1: General characteristics of the sample and comparison between groups with high lean mass and low lean mass

<table>
<thead>
<tr>
<th>Variables</th>
<th>Men (n=37)</th>
<th></th>
<th>P</th>
<th>Women (n=20)</th>
<th></th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>HLM (18)</td>
<td>LLM (19)</td>
<td></td>
<td>HLM (14)</td>
<td>LLM (6)</td>
<td></td>
</tr>
<tr>
<td>Age (years)</td>
<td>68±4±1.0</td>
<td>69±2±7.3</td>
<td>-0.212</td>
<td>62.7±9.5</td>
<td>66.7±10.4</td>
<td>-0.656</td>
</tr>
<tr>
<td>Height (cm)</td>
<td>169±0±3.5</td>
<td>168±0±6.0</td>
<td>0.455</td>
<td>154.5±8.6</td>
<td>154.3±5.1</td>
<td>0.032</td>
</tr>
<tr>
<td>Weight (kg)</td>
<td>80.9±7.9</td>
<td>86±10.8</td>
<td>2.977</td>
<td>61.8±14.3</td>
<td>58.5±3.5</td>
<td>0.386</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>28.3±2.6</td>
<td>24±3.6</td>
<td>2.958</td>
<td>25.8±5.0</td>
<td>24.5±0.5</td>
<td>0.417</td>
</tr>
<tr>
<td>Appendicular lean body mass (kg/m²)</td>
<td>8.4±0.6</td>
<td>7±0.4</td>
<td>7.552</td>
<td>6.3±0.6</td>
<td>5.2±0.1</td>
<td>2.832</td>
</tr>
<tr>
<td>Knee extension (N)</td>
<td>269±6.104.3</td>
<td>199±55.9</td>
<td>2.554</td>
<td>146.3±36.3</td>
<td>124±31.6</td>
<td>0.954</td>
</tr>
<tr>
<td>Knee flexion (N)</td>
<td>118±8.14.2</td>
<td>110±24.1</td>
<td>0.993</td>
<td>75.6±23.1</td>
<td>70.3±5.8</td>
<td>0.388</td>
</tr>
<tr>
<td>Shoulder flexion (N)</td>
<td>63.7±17.5</td>
<td>61.8±21.2</td>
<td>0.235</td>
<td>41.6±12.7</td>
<td>39.7±18.2</td>
<td>0.221</td>
</tr>
<tr>
<td>Shoulder abduction (N)</td>
<td>60±20.0</td>
<td>56.1±18.0</td>
<td>0.639</td>
<td>36.7±11.7</td>
<td>35.7±10.2</td>
<td>0.138</td>
</tr>
</tbody>
</table>

Note: The data are expressed in mean and (±) standard deviation.
Abbreviations: BMI, body mass index; HLM, high lean mass; LLM, low lean mass; N, newton.
Table 2 Association between low lean mass and muscle strength and functional exercise capacity in COPD patients

<table>
<thead>
<tr>
<th>Variables</th>
<th>Low lean mass</th>
<th>OR (CI 95%)</th>
<th>P</th>
</tr>
</thead>
<tbody>
<tr>
<td>Knee extension</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Higher</td>
<td>12 (21.4)</td>
<td>1.00</td>
<td>0.045</td>
</tr>
<tr>
<td>Lower</td>
<td>20 (35.7)</td>
<td>5.93 (1.04–33.74)</td>
<td></td>
</tr>
<tr>
<td>Knee flexion</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Higher</td>
<td>14 (25)</td>
<td>1.00</td>
<td>0.298</td>
</tr>
<tr>
<td>Lower</td>
<td>18 (32.1)</td>
<td>2.20 (0.49–9.68)</td>
<td></td>
</tr>
<tr>
<td>Shoulder flexion</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Higher</td>
<td>16 (28.6)</td>
<td>1.00</td>
<td>0.898</td>
</tr>
<tr>
<td>Lower</td>
<td>16 (28.6)</td>
<td>0.91 (0.20–4.02)</td>
<td></td>
</tr>
<tr>
<td>Shoulder abduction</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Higher</td>
<td>15 (26.8)</td>
<td>1.00</td>
<td>0.848</td>
</tr>
<tr>
<td>Lower</td>
<td>17 (30.3)</td>
<td>0.86 (0.19–3.85)</td>
<td></td>
</tr>
<tr>
<td>Functional exercise capacity</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Higher</td>
<td>14 (25)</td>
<td>1.00</td>
<td>0.112</td>
</tr>
<tr>
<td>Lower</td>
<td>18 (32.1)</td>
<td>0.82 (0.054–1.357)</td>
<td></td>
</tr>
</tbody>
</table>

Abbreviations: OR, odds ratio; CI, confidence interval.

dynamometer. These results are not in line with our findings. We used a fixed dynamometer to evaluate the muscle force while Hillman et al. used a hand-held dynamometer. This difference in the applied methodology may explain the contrasting results.

In our study, muscle mass loss could be inferred by knee-extension strength. A simple, affordable, and widely used measurement in clinical practice provides an idea of muscle characteristics apart from strength. The clinical applicability of our findings is related to an accessible alternative to infer the muscle composition of patients with COPD. This type of evaluation allows the elaboration of a more appropriate therapy and hence, optimizes the treatment of these patients.

A potential limitation of the study is that the sample size was not enough for the results to be extrapolated to the entire population of COPD. Although the establishment of cut-off values of dynamometry to categorize patients into HLM or LLM was not the primary aim of the present study, further research with larger sample would be indicated for this purpose.

In conclusion, despite the small sample, a simple and inexpensive technique proved to be able to infer muscle mass loss in patients with COPD.

Acknowledgments

This work was supported by the following Brazilian Scientific Agencies: Fundação de Amparo à Pesquisa do Estado de São Paulo (FAPESP), Programa Institucional da Pró-Reitoria de Extensão – PROEX/UNESP and FUNDUNESP.

Disclosure

The authors have no financial or nonfinancial conflicts of interest with any company referred to in this article.

References


