CROSS-SECTIONAL PILOT STUDY

Prevalence of sarcopenia among hospitalized internal medicine patients: A cross-sectional single-center pilot study according to EWGSOP2 criteria

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ABSTRACT

INTRODUCTION: Sarcopenia is associated with a higher risk of adverse outcomes among hospitalized patients with chronic diseases. Currently, an unmet need lies in its low diagnostic yield. We conducted a pilot study with the aim to screen hospitalized patients for sarcopenia using the EWGSOP2 criteria. PATIENTS AND METHODS: A cross-sectional study among hospitalized patients in internal medicine was carried out, inclusion criteria: age>55, ability to stand-up from a chair. Exclusion criteria: bed-bound, terminal stages of the disease, ICU stay. We measured hand-grip strength, fat-free mass by bioimpedance and short physical performance battery test (SPPB). Sarcopenia was diagnosed in low hand-grip strength and low muscle mass (EWGSOP2), severe sarcopenia in sarcopenic patients with low physical performance (SPPBT≤8).

RESULTS: 40 patients were enrolled, sarcopenia was diagnosed in 8/20 (40%) men and 3/20 (15%) women (p=0.15), severe sarcopenia in 6/20 men (30%) and 2/20 (10%) women (p=0.24). 65% of men and 40% of women had SPPBT≤8, and 60% of men and 55% of women had gait speed≤0.8m/s.

CONCLUSION: Among hospitalized internal medicine patients sarcopenia and severe sarcopenia are common, particularly in males. New diagnostic criteria provide a relatively simple and applicable tool for screening among internal medicine inpatients (*Tab. 3, Ref. 25*). Text in PDF *www.elis.sk.* KEYWORDS: sarcopenia, prevalence, screening, internal medicine, hospitalized patients.

Introduction

Sarcopenia is defined by a progressive and generalized muscle loss(1). It is most often caused by aging, but it also accompanies chronic diseases such as heart failure(2), renal insufficiency(3), obstructive pulmonary disease(4), and liver cirrhosis(5). It is associated with a higher risk of adverse outcomes such as decreased physical performance, immobility, falls, fractures, repeated hospitalizations, loss of independence on activities of daily living, low quality of life or death(6). The number of elderly people at risk of dependence is increasing while putting pressure on the country's health, social and economic systems(7). One of the unfulfilled goals of current patient management is the lack of diagnosis of sarcopenia(8). Identifying these patients would be helpful since patient management could be adapted to prevent further muscle loss. The prevalence of sarcopenia in healthy individuals aged 60-70 has been reported between 5-13%(9). Among patients older than

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70 hospitalized at internal medicine, the prevalence of sarcopenia has been recently reported as high as 42%(10). With such a high incidence of sarcopenia, it would be appropriate to introduce sarcopenia screening in all hospitalized patients.

Diagnostic criteria for sarcopenia have evolved in recent years from the measurement of muscle mass (EWGSOP) to the recently updated consensus of the European working group for sarcopenia in the elderly people (EWGSOP2)(11). This consensus has provided a simplified approach with an aim to increase the diagnostic yield using muscle strength as a principal criterion for sarcopenia. Therefore, we decided to conduct a pilot study with the aim to screen hospitalized internal medicine patients for sarcopenia using the new consensus criteria.

Aims

To estimate the prevalence of sarcopenia among hospitalized internal medicine patients. First, by comparing the basic anthropometric parameters between the group of hospitalized patients and healthy controls. Second, to estimate the prevalence of sarcopenia and severe sarcopenia among hospitalized patients using the EWGSOP2 consensus criteria. Third, to compare basic anthropometric parameters and serum markers according to sarcopenia status.

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Patients and methods

We conducted a cross-sectional study among patients hospitalized at the 5th Department of Internal Medicine Comenius University Faculty of Medicine during the period of 2 weeks. This pilot study is a preliminary report from a larger study that is being conducted in a prospective manner with an aim of finding the most suitable and informative diagnostic tool/parameter for sarcopenia in our particular case-mix of inpatients. Inclusion criteria were an acute hospitalization, age above 55 years, ability to stand-up from a chair and ability to cooperate in measuring the hand-grip strength. We excluded patients that were bed-bound on admission, patients with terminal stages of any disease, with active malignant disease and those hospitalized at the intensive care unit. The control group comprised of the age and sex-matched community-dwelling healthy individuals in whom we assessed anthropometric measures.

In all patients and controls, we assessed the following anthropometric parameters: body mass index (BMI, kg/m²), mid-arm circumference (cm), tricipital skinfold (cm), subscapular skinfold and the hand-grip strength. Skinfolds were measured using a caliper method (personal body fat tester). Hand-grip strength was assessed using a hand-grip meter (Gima S.p.A. Via Marconi, 1, 20060 Gessate, Italy) by calculating an average value from 3 measurements. Normal hand grip strength values were derived from the EWGSOP2 (men<27 kg, men<16 kg). Patients with values below the cut-off were considered as having low muscle strength.

In the patient group, muscle mass was assessed using an electrical bio-impedance device ImpediMed IMP SF7 (ImpediMed Inc. 5900 Pasteur Court, Suite 125, Carlsbad, CA, USA). The measurement was carried out in a supine position, all the electrodes were placed unilaterally on a non-dominant upper limb (yellow electrode - next to the ulna head, a red electrode at the center of the dorsal arm at a distance of 2-5 cm from the yellow electrode) and lower limb (blue electrode - center between the medial and lateral malleoli, black electrode - the center of the foot dorsum at a distance of 2-5 cm from the blue electrode). From the measured parameters, we assessed the fat-free mass and the fat mass in kg, that was converted into the fat-free mass index (FFMI) and fat mass index (FMI) divided by the square of body height. We adopted normal values for FFMI and FMI from the study among 153 men and 451 women in the geographically and ethnically similar population in Austria(12). Normal values in this study were adjusted according to sex and BMI. Patients with values below the estimated reference range were considered as having a low muscle mass.

Functional status of patients was assessed by the short physical performance battery test(13) (SPPBT) by measuring the following parameters: 1. Chair-stands (s), the time necessary for 5 chair stands without the help of hands, 2. balance test (s), time the patient maintains the balance having feet together, at the semitandem and tandem position for maximum 10 seconds in each. 3. Gait speed, by measuring the time necessary to walk 5 meters (m/s). From the measured parameters, we calculated the SPPBT score. Patients having an SBBPT score ≤ 8 were considered as having a low physical performance.

Sarcopenia was diagnosed according to the criteria of the European working group for sarcopenia in elderly people (EWGSOP2) in patients having low hand-grip strength and a low muscle mass (11). Severe sarcopenia was diagnosed in patients having sarcopenia and low physical performance.

We also assessed basic laboratory parameters (hemoglobin, serum creatinine, CRP, serum vitamin D) for comparison between sarcopenic and non-sarcopenic patients.

The study was conducted in accordance with the proceeding of the declaration of Helsinki. All participants signed an informed consent prior to any procedure. The study protocol was approved by the local ethics committee.

Statistical analysis was carried out using R statistical software version 3.5.1 (R foundation for statistical computing, Vi-

Tab. 1. Com	parison of p	arameters	between 1	the study	group and	control s	group.

		Healthy controls	Patients	*	
		Women, n (%), median $[IQR]$ n = 20	n = 20	- p*	
Age category (%)	56 to 70	10 (50.0)	12 (60.0)	- 0.751	
	71 to 85	10 (50.0)	8 (40.0)		
Body mass index (kg/m ²)		25.73 [23.43, 27.34]	27.27 [25.35, 32.09]	0.062	
Tricipital skinfold (cm)		1.6 [1.28, 1.92]	2.5 [2.0, 2.92]	< 0.001	
Mid-arm circumference (cm)		28.0 [27.0, 31.0]	31.0 [27.5, 33.0]	0.149	
Hand-grip strength (kg)		19.41 [17.04, 21.25]	16.83 [14.58, 21.08]	0.387	
Hand-grip strength <16kg		4 (20.0)	8 (40.0)	0,173	
		Men, n (%), median [IQR] n = 20	n = 20		
A (0/)	56 to 70	10 (50.0)	11 (55.0)	1	
Age category (%)	71 to 85	10 (50.0)	9 (45.0)	- 1	
Body mass index (kg/m ²)		28.1 [25.45, 29.81]	27.4 [26.0, 29.04]	0.957	
Tricipital skinfold (cm)		1.75 [1.50, 1.92]	1.40 [1.08, 2.05]	0.261	
Mid-arm circumference (cm)		32.0 [28.75, 34.25]	30.0 [28.0, 31.25]	0.128	
Hand-grip strength (kg)		31.9 [24.58, 37.67]	25.5 [19.1, 29.83]	0.024	
Hand-grip strength<27kg		6 (30.0)	12 (60.0)	0.06	

*difference between patients and healthy controls, chi-square or Fisher test, or Mann-Whitney test

enna Austria) using an EZR Plugin (Kanda et al. (14)) Results of numerical variables are displayed as medians and interquartile ranges, proportions are displayed as numbers and percentages. Mann-Whitney test was used for comparison of numerical variables and the chi-square test or Fisher test was used to compare proportions. Significant differences were stated in cases when the probability of null-hypothesis was inferior to 5% (p<0.05).

Results

We enrolled 40 patients, 20 men and 20 women, and 40 healthy controls, 20 men and 20 women.

Sex-specific comparison for anthropometric parameters between the patient group and the control group are displayed in Table 1. In women, patients had significantly higher tricipital skinfold compared to healthy controls (2.5 IQR2.0-2.92 vs. 1.6 IQR1.28-1.92, p<0.001) and a trend towards higher BMI (27.27 IQR 25.35-32.1 vs. 25.73 IQR23.43-27.34, p=0.062), but there was no difference in the hand-grip strength (16.83 IQR14.58-21.08 vs. 19.41IQR 17.04-21.25, p=0.39) or the prevalence of HGS below 16kg. In men, patients had comparable BMI and the tricipital skinfold but had lower hand-grip strength and a higher prevalence of hand-grip strength below 27 kg compared to healthy controls

Tab. 2. Summary statistics and characteristics of the patient group

		Men, n = 20	Women, $n = 20$	- P*	
		Median [IQR], n (%)	Median [IQR], n (%)	- P*	
Age		70.00 [60.75, 74.75]	69.50 [65.00, 73.00]	0.90	
BMI (kg/m ²)		27.38 [26.00, 29.04]	27.27 [25.35, 32.09]	0.93	
Principal diagnosis					
Dg. Anemia		2 (10.0)	3 (15.0)		
Dg. Gastrointestinal disease		2 (10.0)	4 (20.0)		
Dg. Renal disease		1 (5.0)	1 (5.0)	0.86	
Dg. Cardiovascular disease		9 (45.0)	7 (35.0)	0.80	
Dg. Metabolic disease		3 (15.0)	4 (20.0)		
Dg. Sepsis		3 (15.0)	1 (5.0)		
Waist circumference (cm)		106.50 [101.75, 113.25]	100.50 [91.75, 114.50]	0.33	
Mid-arm circumference (cm)		30.00 [28.00, 31.25]	31.00 [27.50, 33.00]	0.56	
Subscapular skinfold (cm)		2.00 [1.87, 2.23]	2.05 [1.80, 2.70]	0.61	
Triceps skinfold (cm)		1.40 [1.08, 2.05]	2.50 [2.00, 2.92]	<0.01	
Hand-grip strength (kg)		25.50 [19.08, 29.83]	16.83 [14.58, 21.08]	0.01	
Low hand-grip strength	no	8 (40.0)	12 (60.0)	0.04	
(men<27, women<16 kg)	yes	12 (60.0)	8 (40.0)	- 0.34	
Balance test (s)		30.00 [23.50, 30.00]	30.00 [27.25, 30.00]	0.48	
Gait-speed (m/s)		0.60 [0.42, 1.05]	0.71 [0.52, 1.10]	0.24	
	no	8 (40.0)	9 (45.0)	- 1.00	
Low gait speed (≤0.8 m/s)	yes	12 (60.0)	11 (55.0)		
Five chair stands (s)	<u>y</u> es	14.55 [11.00, 23.10]	13.25 [11.95, 15.57]	0.56	
Tive chair stands (3)	no	12 (60.0)	14 (70.0)	0.74	
Chairstands>15s		8 (40.0)	6 (30.0)	- 0.74	
SPPBT <i>f</i> score	yes	6.00 [5.75, 11.00]	9.50 [6.75, 10.25]	0.41	
SPFB1/ scole				0.41	
SPPBT score ≤8	no	7 (35.0)	12 (60.0)	0.21	
	yes	13 (65.0)	8 (40.0)		
Fat-free mass index (kg/m ²)		20.59 [19.03, 22.35]	17.76 [16.18, 20.89]	0.04	
Fat-free mas index < normal	no	8 (40.0)	14 (70.0)	0.11	
	yes	12 (60.0)	6 (30.0)		
Fat mass index (kg/m ²)		7.06 [4.98, 10.17]	9.87 [6.79, 11.49]	0.22	
Fat mass index < normal	no	19 (95.0)	15 (75.0)	0.18	
r'at mass muex < norman	yes	1 (5.0)	5 (25.0)	0.18	
Serum kreatinin (umol/l)		90.70 [76.77, 124.25]	73.35 [60.95, 110.10]	0.13	
Hemoglobin (g/l)		122.50 [110.3, 131.3]	120.00 [110.0, 128.3]	0.77	
C-reactive protein (mg/l)		6.15 [1.15-47.56]	4.43 [1.82, 26.97]	0.86	
Serum vitamin D (ug/l)		16.95 [12.15, 22.00]	20.05 [14.73, 27.00]	0.32	
	no	12 (60.0)	17 (85.0)		
Sarcopenia	yes	8 (40.0)	3 (15.0)	- 0.16	
	no	14 (70.0)	18 (90.0)	- 0.24	
Severe sarcopenia		6 (30.0)	2 (10.0)		
	yes	0 (30.0)	2 (10.0)		

* p value for the sex difference, chi-square or Fisher test or Mann-Whitney test

fSPPBT, short performance physical battery test

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Tab. 3. Comparison of patients' characteristics according to the presence of sarcopenia.

	No sarcopenia, $n = 29$	Sarcopenia, n = 11	- p/	
	Median [IQR], n (%)	Median [IQR], n (%)	- P'	
Age	69.00 [64.00, 73.00]	71.00 [64.00, 75.50]	0.504	
Age category (56-70)	18 (62.1)	5 (45.5)	- 0.477	
Age category (71-85)	11 (37.9)	6 (54.5)		
Men	12 (41.4)	8 (72.7)	- 0.155	
Women	17 (58.6)	3 (27.3)		
Height (cm)	170.00 [164.00, 178.00]	166.00 [165.00, 172.50]	0.585	
Weight (kg)	83.00 [74.00, 95.00]	74.00 [66.00, 79.00]	0.108	
BMI (kg/m ²)	27.77 [25.62, 31.84]	26.90 [24.58, 28.78]	0.303	
BMI category (18-24.9)	8 (27.6)	3 (27.3)		
BMI category 25-29.9)	13 (44.8)	6 (54.5)	0.9	
BMI category >30	8 (27.6)	2 (18.2)	-	
Principal diagnosis				
Dg. Anemia	4 (13.8)	1 (9.1)		
Dg. Gastrointestinal disease	5 (17.2)	1 (9.1)	0.932	
Dg. Renal disease	2 (6.9)	0 (0.0)		
Dg. Cardiovascular disease	11 (37.9)	5 (45.5)	0.932	
Dg. Metabolic disease	4 (13.8)	3 (27.3)		
Dg. Sepsis	3 (10.3)	1 (9.1)		
Waist circumference (cm)	108.00 [99.00, 115.00]	106.00 [98.00, 110.00]	0.534	
Mid-arm circumference (cm)	31.00 [29.00, 33.00]	29.00 [26.50, 30.50]	0.038	
Subscapular skinfold (cm)	2.10 [1.80, 2.50]	2.00 [1.85, 2.30]	0.419	
Triceps skinfold (cm)	2.30 [1.50, 2.60]	1.50 [1.15, 2.35]	0.288	
Balance test (s)	30.00 [25.00, 30.00]	30.00 [23.00, 30.00]	0.538	
Gait-speed (m/s)	0.71 [0.50, 1.00]	0.50 [0.33, 1.22]	0.275	
Five chair stands (s)	13.00 [11.00, 15.50]	18.00 [12.75, 24.52]	0.066	
Fat mass index (kg/m ²)	8.50 [6.11, 10.11]	9.53 [6.48, 11.21]	0,617	
Serum bilirubin (umol/l)	10.90 [9.17, 15.00]	12.00 [9.65, 19.59]	0.535	
Serum kreatinin (umol/l)	85.10 [70.80, 114.00]	87.10 [66.45, 149.90]	0.515	
C-reactive protein (mg/l)	6.3 [1.97, 29.47]	3.4 [2.02, 22.31]	0.75	
Serum urea (mmol/l)	5.84 [4.73, 8.68]	5.10 [4.01, 9.63]	0.94	
Hemoglobin (g/l)	120.00 [107.00, 128.00]	123.00 [116.50, 137.00]	0.296	
Serum vitamin D (ug/l)	18.40 [14.10, 25.20]	18.00 [14.50, 22.10]	0.976	

^fchi-square or Fisher test, or Mann-Whitney test

(25.5 IQR19.1-29.83 vs. 31.9 IQR24.58-37.67, p=0.024 and 60% vs. 30%, p=0.06).

Among 40 hospitalized patients, the median age in both sexes was 70.0 vs. 69.5 respectively. Study group characteristics and the comparison between men and women are displayed in Table 2. Median BMI was in the range of overweight (27.38 kg/m² and 27.27, p=0.93). Women had significantly higher tricipital skinfold compared to men (2.5 cm IQR2.0-2.92 vs. 1.4 IQR1.08-2.05), lower hand-grip strength (16.83 kg IQR14.58-21.08 vs. 25.5 IQR19.08-29.83, p=0.01) and lower fat-free mass index (17.76 IQR16.18-20.89 vs. 20.59 IQR 19.03-22.35, p=0.04). Among the parameters of functional status (chair stands, gait speed, balance) we did not observe any difference between the sexes.

Sarcopenia was diagnosed in 8/20 (40%) men and 3/20 (15%) women (p=0.15). Severe sarcopenia was diagnosed in 6/20 men (30%) and 2/20 (10%) women (p=0.24). According to the SPPB test, 65% of men and 40% of women had low performance status. According to the gait speed, 60% of men and 55% of women had low gait speed. The differences were not statistically significant.

Comparison of selected parameters, with the exception of those that defined sarcopenia, between the group of patients with and without sarcopenia is displayed in Table 3. We observed no difference in the serum parameters between the groups. Serum vitamin D concentrations were not significantly different between patients with or without sarcopenia, between patients with low or normal hand grip strength, gait speed or SPPB score. Sarcopenic patients had lower mid-arm circumference (29.0 IQR26.5-30.5 vs. 31.0 IQR29.0-33.0, p=0.038) and a trend towards higher time for chairstands (18.0 IQR12.75-24.52 vs. 13.0 IQR11.0-15.5, p=0.06).

Discussion

In our study, we aimed at sarcopenia screening among hospitalized internal medicine patients. Hospitalized women compared to healthy controls had significantly higher tricipital skinfold while hospitalized men had significantly lower hand grip strength. Among 40 hospitalized patients using the EWGSOP2 criteria, sarcopenia was diagnosed in 40% of men and 15 % of women, severe sarcopenia in 30% of men and 10% of women. Patients with or without sarcopenia did not differ in basic laboratory parameters including vitamin D.

This pilot study is an inaugural report with an obvious limitation allowing only for cross-sectional analysis. Clinical significance of our findings and their impact on the outcome of patients will be tested in a prospective manner and by including a higher number of patients. However, this pilot study confirms that the new consensus criteria for diagnosing sarcopenia are feasible in our clinical practice. Hand-grip strength measurement is an easy screening tool that is objective and requires little time for evaluation. Furthermore, bioimpedance techniques for evaluating the muscle mass could be performed bed-side with little dedicated time. These two tests appear as the simplest choice since other alternatives proposed by the current consensus (DXA scan, or CT/ MRI scans) are more time and device demanding. For confirming severe sarcopenia, physical performance tests are more time-consuming. We decided to use a more complex SPPBT, but the current consensus also proposes using the gait speed. We also tested gait speed, and the difference in the diagnostic yield for severe sarcopenia would only be in one male patient decreasing its prevalence from 20 to 17.5%. In clinical practice, there is an advantage in physical testing because it could be repeatedly performed during hospitalization for evaluating early changes in physical status. Some prospective studies have reported, that in a particular group of patients, not all physical performance parameters have an equal impact on the outcome. For example, a study among very old men from Sweden, hand-grip strength and gait speed were associated with independent aging, but not muscle mass or diagnosed sarcopenia(15). Only the prospective arm of our study will find the most informative parameter(s) in the prediction of the adverse outcome among our particular case-mix of inpatients.

The prevalence of sarcopenia and severe sarcopenia in 40% or 30% of men and 15% and 10% of women was high and suggests that sarcopenia should be screened in patients hospitalized at internal medicine. Some previous reports have found similar prevalence among hospitalized medical patients (16). Among hospitalide geriatric patients, the prevalence of sarcopenia ranged from 11.4% (17) up to 51% and 64% among acute geriatric patients or elderly patients after myocardial infarction (18,19). However, the previous studies have used different diagnostic criteria not allowing for a direct comparison with our study using EWGSOP2 criteria. One study has compared the prevalence of sarcopenia using both EWGSOP1 and 2 algorithms showing a slightly lower prevalence of 27.7 vs.18.1% when using EWGSOP2 criteria (20).

The presence of sarcopenia among hospitalized patients increases the risk of adverse outcome. During 2 years after an acute hospitalization, the risk of death has been reported as 4 to 9 times higher (17,18). A systematic review has also reported that sarcopenia increases the risk of hospital related outcomes in elderly people by more than 40% (21). Therefore, due to the risk of adverse outcome during hospitalization, physicians have an opportunity for applying early measures for preventing further loss of muscle mass and function. Currently, early nutritional intervention with increased protein consumption, supplementation of vitamin D as well as regular physiotherapy have been recommended by the recent guidelines(22,23). Finally, we did not find any significant difference in the vitamin D levels according to sarcopenia status, low hand-grip strength, low gait speed or low physical performance, but this pilot study was not powered for detecting a difference in laboratory parameters.

In our study, the prevalence of sarcopenia and severe sarcopenia was higher in men than in women. The difference was not statistically significant but this was likely due to low numbers (p=0.15). The prevalence of low functional status was also numerically higher in men (65%) than in women (40%), but low gait speed was found among similar proportions of patients (60% vs. 55%). Some sex-related differences in the prevalence of sarcopenia and its outcome have been reported (19,20,24). It appears that in men, chronic diseases lead to a more dynamic decline in muscle mass. In contrast, due to different fat distribution, hospitalized women had significantly higher tricipital skin-fold and also higher BMI compared to healthy controls, probably reflecting the greater burden of the metabolic syndrome and its complications on women. This finding also points to an important sex-related difference in body composition and changes in body composition in chronic disease. One recent study has shown that in women with cirrhosis, it was the subcutaneous fat loss and not the muscle mass that was associated with the adverse outcome (25). Therefore, the prognostic value of sarcopenia and functional status should always be separately reported for each sex.

Conclusion

Among hospitalized internal medicine patients by applying the updated EWGSOP2 criteria we diagnosed sarcopenia and severe sarcopenia in 40% and 30% of men and 15% and 10% of women. The new diagnostic criteria were confirmed as easily applicable for inpatients. Observed high prevalence of sarcopenia justifies its screening among all internal medicine inpatients allowing for early preventive or therapeutic interventions.

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