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4 **Can early assessment of hand grip strength in elderly hip fracture patients predict**
5 **functional outcome?**

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23 **Abstract**

24 Decreased muscle strength is not only a risk factor for hip fracture in elderly patients, but plays a role in recovery of
25 physical function. Our aim was to assess the role of grip strength measured early after hip fracture, and classified according to the
26 EWGSOP2 criteria in predicting short- and long-term functional recovery. One hundred ninety-one patients with acute hip fracture
27 consecutively admitted to an orthopaedic hospital have been selected. A multidimensional geriatric assessment evaluating
28 sociodemographic variables, cognitive status, functional status and quality of life prior to fracture, as well as perioperative variables
29 were performed. Follow-ups at 3 and 6 months after surgery were carried out to evaluate functional recovery. Multivariate
30 regression models were used to assess the predictive role of handgrip strength. The mean age of the participants was 80.3 ± 6.8
31 years. Thirty-five percent of our patients with clinically relevant hand grip strength weakness were significantly older, more often
32 female, had a lower BMI, and were of worse physical health. They also had a lower cognitive level, lower Barthel index, and lower
33 EQ5D scores before fracture. Multivariate regression analysis adjusted for age and gender revealed that hand grip weakness was an
34 independent predictor of worse functional outcome at 3 and 6 months after hip fracture for both genders and in all age populations.
35 Our study supports the prognostic role of hand grip strength assessed at hospital admission in patients with hip fracture. Thus,
36 clinicians should be encouraged to include hand grip assessment in their evaluation of hip fracture patients in the acute setting in
37 order to optimize treatment of high-risk individuals.

38 **Key words:** hip fracture, hand grip strength, functional recovery, sarcopenia, frailty

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40 **Introduction**

41 Sustaining a hip fracture is considered one of the most fatal fractures for elderly people that leads to impaired function, and
42 increased morbidity and mortality, and high financial liability. These facts challenge clinicians in identifying patients at risk of
43 worse outcome early in the course of hip fracture treatment, in order to set realistic rehabilitation goals, optimize perioperative care,
44 and define optimal rehabilitation strategies in order to reduce devastating outcomes.

45 Functional evaluation in patients with hip fracture is an essential part of multidimensional assessment, and has an important
46 prognostic value. Muscle weakness is considered a key element of frailty [1] and, increasingly, of sarcopenia [2, 3]. It is believed
47 that sarcopenia not only enhances fracture risk, but also increases the risk of poor functional outcome after hip fracture [4].
48 Reduced muscle strength makes it more difficult to regain lost balance and decreases the mechanical loading of the skeleton leading
49 to reduced adaptive bone remodeling [5, 6]. Hand grip strength (HGS) assessment is an objective measure of overall body muscle
50 strength and physical function [7, 8], an important measure for frailty [9], and sarcopenia [10, 11]. Various studies have shown the
51 prognostic value of hand grip strength in patients with hip fracture [12-17]. However, very few have been carried out in the acute
52 phase [16, 17], but there are no studies which have been carried out using the European Working Group on Sarcopenia in Older
53 People 2 (EWGSOP2) criteria [3] to define clinically relevant hand grip weakness.

54 The aim of our study was to assess the EWGSOP2 threshold for grip strength assessed at admission to hospital after hip fracture to
55 predict short- and long-term functional recovery. We hypothesized that levels of grip strength below the EWGSOP 2 thresholds
56 measured in the first 48h after hip fracture could predict an unfavorable short- and long-term functional outcome.

57

58 **Materials and methods**

59 **Study design**

60 All adult patients 65 years or older with an acute hip fracture who were admitted consecutively to an university associated
61 orthopedic hospital in Serbia between March 1st 2017 and February 28th 2018 were enrolled in an open, prospective, observational
62 cohort study. All patients with pathologic fractures, major concomitant injuries, multiple trauma, malignant diseases, imminent
63 death as a result of an end-stage disease, inability to walk before fracture, and nonoperative treatment resulting from high surgical
64 risk were excluded. Furthermore, patients with severe cognitive impairment, as well as patients with hand weakness as a
65 consequence of previous neurologic disorders or hand injuries also were excluded. During the study period, 551 patients had hip
66 fractures and were examined for eligibility. One hundred ninety-one patients were confirmed eligible and were included in the
67 study. All patients gave written informed consent to participate in the study. The study was conducted according to the Helsinki
68 Declaration and approved by the University's institutional review board.

69 **Measures**

70 **Baseline evaluation**

71 We assessed all subjects through standardized patient interview with respect to sociodemographic variables (age, sex,
72 marital status, preinjury living conditions), cognitive level, handgrip strength, prefracture functional level, and health related quality
73 of life within 24h of admission. We also recorded perioperative variables during the primary hospital stay, such as comorbidity
74 level, waiting time for surgery, type of fracture, surgical method, type of anesthesia, and presence of postoperative complications,
75 and length of stay (LOS).

76 Cognitive level was assessed with the Short Portable Mental Status Questionnaire (SPMSQ) [18]. The 10-item questionnaire
77 classifies the patient's cognitive level depending on the number of correct answers as lucid (8–10), mild to moderate cognitive
78 dysfunction (3–7), and severe cognitive dysfunction (0–2). Handgrip strength was measured using a JAMAR hand dynamometer
79 (Model BK-7498, Fed Sammons Inc, Brookfield, III). Patients were in the supine position, and encouraged to exhibit the greatest
80 possible force [19]. The best recorded of 3 attempts of maximal voluntary contraction performed at 1-minute intervals of the
81 dominant hand was considered for analysis. Hand grip strength measurements less than 16 kg in women and 27 kg in men were
82 considered cut-points for the diagnosis of sarcopenia according to the revised EWGSOP2 criteria [3]. The pre-fracture functional
83 status 2 weeks before hospital admission was assessed by the Barthel index [20]. The Barthel index measures performance in basic
84 activities of daily living; its score ranges from 0 (total dependence) to 100 (total independence) [21]. General health related quality

85 of life was measured with the EQ5D scale, which consists of a five-level response for five domains related to daily activities,
86 mobility, self-care, usual activities, pain and discomfort, anxiety and depression [22]. Responses to the health status classification
87 system are converted into an overall score using a published utility algorithm for the UK population [23].

88 We used the Charlson comorbidity index (CCI) to categorize comorbidities [24]. Patients were divided into three groups:
89 without and mild, with CCI scores of 1–2; moderate, with CCI scores of 3–4; and severe, with CCI scores ≥ 5 .

90 All patients with femoral neck fractures (84 patients (43.9%)) underwent bipolar hemiarthroplasty, whereas all patients with
91 intertrochanteric (92 patients (48.2%)) and subtrochanteric fractures (15 patients (7.9%)) underwent open reduction and internal
92 fixation (ORIF). In all patient early assisted ambulation was encouraged on the first postoperative day with weightbearing as
93 tolerated, and all patients followed a standardized postoperative rehabilitation program.

94 **Outcomes**

95 Functional status after 3 and 6 months was evaluated using the Barthel index score. The information was collected by phone
96 interview. Data from patients who died or were lost before the first and second follow-up respectively were excluded from the
97 study. For the analysis of Barthel index 3 months postoperatively the sample size included 160 patient (22 (11.5%) died; 9 (4.7%)
98 were lost to follow-up). Analysis of outcomes six months after the fracture was performed on 154 patients (27 (14.1%) died, 10
99 (5.3%) were lost to follow-up).

100 **Statistical analysis**

101 Continuous variables are presented in terms of mean values with SD or median and interquartile range depending on
102 Kolmogorov-Smirnov test of distribution normality. Categorical values are summarized as absolute frequencies and percentages.
103 To compare patients with two different categories of grip strength a *t* test was performed for the continuous variables and a Mann-
104 Whitney U test for ordinal variables.

105 In order to detect potential and independent predictors of recovery expressed as Barthel index scores after 3 and 6 months,
106 univariate and then multivariate linear regression with collinearity diagnostic (VIF method used; variables with VIF > 4 were
107 excluded from multivariate models) was used. Both multivariate models were adjusted for age and gender.

108 The significance level for all statistical tests was set at 0.05. All analyses were performed using the SPSS Inc. Released
109 2008. SPSS Statistics for Windows, Version 17.0. Chicago: SPSS Inc.

110

111 **Results**

112 Our cohort consisted of 191 patients aged 66 to 97 years. The mean age was 80.3 ±6.8 years, and 77.0% of our cohort were
113 women. The mean HGS in our cohort was 20.5 ±6.8 kg. Sixty-six (34.6%) patients had clinically relevant hand grip weakness.
114 Those patients were significantly older, more often female, had a lower BMI, and were of worse physical health. They also had a

115 lower cognitive level, lower Barthel index scores and lower EQ5D scores before fracture. Patients with weaker grip strength were
 116 more often operated in general anesthesia (Table 1).

117

118 **TABLE 1. Socio-demographic and baseline pre- and perioperative characteristics of the participants**

	Women With HGS < 16 kg Men With HGS < 27 kg N=66 (34.6%)	Women With HGS ≥ 16 kg Men With HGS ≥ 27kg N=125 (65.4%)	p
Age (year)*	83.53 ± 6.16	78.52 ± 6.50	<0.001
Gender**			
Male	21 (31.8%)	23 (18.4%)	0.036
Female	45 (68.2%)	102 (81.6%)	
Marital status**			
Other	40 (61.5%)	76 (61.8%)	0.973

Married	25 (38.5%)	47 (38.2%)	
Pre-injury residence**			
Home (live alone)	14 (21.2%)	31 (24.8%)	0.710
Home (live with family)	50 (75.8%)	92 (73.6%)	
Institution	2 (3.0%)	2 (1.6%)	
BMI*	23.82 ± 4.77	25.60 ± 3.91	0.008
CCI groups**			
No comorbidity/mild	21 (31.8%)	68 (54.4%)	0.009
Moderate	36 (54.6%)	42 (33.6%)	
Severe	9 (13.6%)	15 (12.0%)	
SPMSQ*	6.79 ± 1.67	7.83 ± 1.63	<0.001
EQ5D before fracture*	0.73 ± 0.17	0.83 ± 0.14	<0.001

Barthel index before fracture*	92.95 ± 7.70	96.16 ± 5.54	0.003
Type of fracture**			
Femoral neck	25 (37.9%)	59 (47.2%)	0.462
Intertrochanteric	35 (53.0%)	57 (45.6%)	
Subtrochanteric	6 (9.1%)	9 (7.2%)	
Time from admission to operation*	6.26 ± 3.17	5.75 ± 2.98	0.277
Lenght of hospital stay*	15.91 ± 5.20	16.03 ± 4.42	0.864
Surgical procedure**			
Arthroplasty	27 (36.4%)	57 (45.6%)	0.219

ORIF	39 (63.6%)	68 (54.4%)	
Type of anaesthesia**			
General	51 (79.7%)	78 (64.5%)	0.032
Regional	13 (20.3%)	43 (35.5%)	
Duration of anaesthesia*	118.47 ± 39.54	114.70 ± 27.18	0.504
Complications**			
Yes	18 (27.3%)	30 (24.0%)	0.620
No	48 (72.7%)	95 (76.0%)	

119 *Values are given as the mean with the standard deviation in parentheses

120 ** Values are given as the number of patients with the percentage in parentheses

121 RR - relative risk; BMI - body mass index; CCI - Charlson Comorbidity Index; SPMSQ - Short Portable Mental Status

122 Questionnaire; HGS - Handgrip strength

123

124 Patients with relevant hand grip weakness achieved statistically significant lower Barthel index scores 3 (56.30 \pm 25.87 vs.
125 75.77 \pm 21.49) (Table 2) and 6 months (67.77 \pm 29.15 vs. 87.66 \pm 19.30) (Table 3) after hip fracture. Adjusted multivariate regression
126 analysis revealed that hand grip strength below the cutoff point for sarcopenia according to the EWGSOP2 was an independent
127 predictor of worse functional outcome at 3 and 6 months after hip fracture for both genders and in all age populations.

128

129 **TABLE 2. Univariate and Multivariate analysis for variables significantly associated with Barthel index scores 3 months**
130 **after fracture**

Predictors	Univariate analysis		Multivariate analysis	
	B (95% CI)	p value	B (95% CI)	p value
Marital status	-0.05 (-11.04 – 5.71)	0.531		
Preinjury residence	-0.18 (-17.22 - -1.41)	0.021	-0.13 (-13.29 - -0.34)	0.039
BMI	0.02 (-0.79 – 1.08)	0.761		
CCI	-0.29 (-15.40 - -5.24)	<0.001	-0.10 (-8.12 – 1.06)	0.131

SPMSQ	0.27 (1.72 – 5.97)	<0.001	0.72 (-0.92 – 2.94)	0.302
EQ5D before fracture	0.31 (25.94 – 71.55)	<0.001	0.16 (2.10 – 46.10)	0.032
Barthel index before fracture	0.51 (1.37 – 2.30)	<0.001	0.36 (0.77 – 1.71)	<0.001
HGS	0.31 (8.29 – 24.51)	<0.001	0.185 (2.29 – 16.71)	0.010
Time from admission to operation	-0.01 (-1.36 – 1.13)	0.860		
Lenght of hospital stay	-0.04 (-1.04 – 0.62)	0.616		
Type of anaesthesia	0.09 (-3.22 – 12.55)	0.244		
Duration of anaesthesia	-0.21 (-0.31 – 0.05)	0.009	-0.12 (-0.20 – 0.01)	0.069
Complications	-0.16 (-18.58 - -0.44)	0.040	-0.13 (-14.90 - -0.20)	0.044

131 Adjusted for age and gender

132 BMI - body mass index; CCI - Charlson Comorbidity Index; SPMSQ - Short Portable Mental Status Questionnaire; HGS -

133 Handgrip strength

134

135 **TABLE 3. Univariate and Multivariate analysis for variables significantly associated with Barthel index scores 6 months**
 136 **after fracture**

Predictors	Univariate analysis		Multivariate analysis	
	B (95% CI)	p value	B RR (95% CI)	p value
Marital status	-0.12 (-14.75 – 2.56)	0.166		
Preinjury residence	-0.14 (-15.52 – 1.20)	0.093		
BMI	0.04 (-0.75 – 1.28)	0.602		
CCI	-0.31 (-16.61 - -5.58)	<0.001	-0.17 (-10.99 - -1.03)	0.018
SPMSQ	0.29 (1.84 – 6.30)	<0.001	0.09 (-0.76 – 3.26)	0.222
EQ5D before fracture	0.36 (32.11 – 78.99)	<0.001	0.15 (-0.28 – 45.56)	0.053
Barthel index before fracture	0.53 (1.47 – 2.48)	<0.001	0.38 (0.86 – 1.91)	<0.001

HGS	0.36 (10.39 – 27.18)	<0.001	0.21 (3.07 – 18.37)	0.006
Time from admission to operation	-0.07 (-1.87 – 0.74)	0.392		
Length of hospital stay	0.01 (-0.83 – 0.94)	0.895		
Type of anaesthesia	0.08 (-4.19 – 12.28)	0.333		
Duration of anaesthesia	-0.17 (-0.28 – -0.01)	0.037	-0.08 (-0.17 – 0.05)	0.250
Complications	-0.07 (-14.06 – 5.82)	0.414		

137 Adjusted for age and gender

138 BMI - body mass index; CCI - Charlson Comorbidity Index; SPMSQ - Short Portable Mental Status Questionnaire; HGS -

139 Handgrip strength

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141 Besides hand grip strength. living at home, better quality of life and higher functionally independence before fracture, as

142 well as absence of complications during the hospitalization period were independent predictors of Barthel index scores 3 months

143 postoperatively (Table 2). Multivariate regression analysis showed that, besides hand grip strength above cutoff values for
144 sarcopenia, lower CCI index and higher Barthel index scores before fracture were independent predictors of higher Barthel index
145 scores 6 months after fracture (Table 3).

146

147 **Discussion**

148 Our study demonstrates that hip fracture patients with a validated threshold for clinical weak grip strength assessed at an
149 early stage had significantly poorer functional recovery after 3 and 6 months compared to patients with a grip strength above the
150 cutoff points. Furthermore, our findings provide evidence that hand grip strength along with several other prognostic factors
151 traditionally considered in clinical practice can independently predict short- and long-term functional outcome [25]. Within the
152 study 35% of the study population had relevant clinical weakness based on hand grip strength. Older age, higher level of
153 comorbidity, lower cognitive level, lower functional level, and worse quality of life recognized at admission indicate clearly a
154 decline of reserve and function across multiple physiological systems in this group of patients.

155 The results of our investigation are consistent with data from previous studies confirming the prognostic role of handgrip
156 strength [12-17]. However, it is not easy to compare present results with other studies addressing this subject. First, handgrip
157 strength was assessed at various time points in different studies. There are only two studies evaluating the prognostic value of
158 handgrip strength measured in the acute setting [16, 17]. Savino et al. showed that handgrip strength measured at hospital admission

159 significantly predicted walking recovery 12 months after hip [16]. Alvarez MN et al. concluded that HGS assessed in the first hours
160 after hospital admission for hip fracture surgery is an indicator of functional recovery after three months [17]. It is well known that
161 muscle mass is maintained during the first 10 days, although it subsequently diminishes [26, 27]. Consequently, it is reasonable to
162 assume that measuring HGS early after hip fracture is an appropriate time to assess function. Second, a small body of literature used
163 cutoff points to define clinical relevant weakness based on HGS [15, 17, 28], and no study applied the EWGSOP2 criteria. Alvarez
164 MN et al. [17] who used the EWGSOP criteria [11], and di Monaco et al. [15] who used the FNIH Sarcopenia Project criteria for
165 HGS categorization [29] confirmed the prognostic role of hand grip strength. In contrast, Steihaug et al. who applied the EWGSOP
166 criteria to investigate the impact of HGS early after fracture were the only ones who found no association between grip strength and
167 short- and long-term functional outcome [28]. This is the only study to deny the value of HGS in predicting functional outcome in
168 hip fracture patients. It has to be taken into account that although the definition of „weakness“ according to the FNIH , and the
169 EWGSOP criteria correspond very closely to the one defined by the EWGSOP2 [3], it is not the same. Moreover, our results
170 cannot be compared completely to those published by de Monaco et al., because they reported their results only on women, and in
171 the postacute rehabilitation setting [13, 15].

172 There are several strengths of our study. First, to the best of our knowledge this is the first study to assess HGS using the
173 EWGSOP2 criteria. Second, our study proves the prognostic value of HGS in the acute setting for both gender and all ages. Most
174 studies who analyzed the predictive role of hand grip strength reported their results only in women with hip fracture [12, 13, 15].

175 There are also some limitations to our study. First, the outcome of our study was assessed with only self-reported information
176 collected by phone interviews. Second, patients were collected only from one single center. Additionally, there are other
177 confounding factors that could have been studied, for example nutritional status and vitamin D status.

178 Our results have two clinical implications. First, assessment of HGS can be used to identify hip fracture patients at high risk
179 for poor functional outcome at an early time point. Second, it is well known that functional evaluation is hard to assess in acute hip
180 fracture patients. In the first place, gait speed cannot be assessed before surgery. Thus, functional evaluation is limited to measuring
181 muscle strength. Confirmation of the prognostic value of HGS assessed in the acute setting is therefore very significant. Third,
182 muscle weakness is a modifiable risk factor that can be improved. It is well known that strengthening exercises had favorable effects
183 on various outcomes after hip fracture [30]. Therefore, future studies should reveal if patients with clinically defined weakness who
184 sustain a hip fracture could benefit from interventions to improve muscle strength, function, and outcomes.

185 Our study has identified HGS assessed in the acute setting as potential prognostic predictor of functional outcome in
186 patients with hip fracture. Hand grip strength is an accessible, cost effective, and simple objective measure of physical function for
187 bedridden patients. Thus, clinicians should be encouraged to include hand grip assessment in their evaluation of hip fracture
188 patients at admission to the acute setting in order to optimize prognostic counseling and treatment of high-risk individuals. Further
189 studies are needed to investigate the relevance of early introduction of resisting exercise programs in patients hip fracture patients
190 with relevant low muscle strength.

191

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