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# Malnutrition in postacute geriatric care: Basic ESPEN diagnosis and etiology based diagnoses analyzed by length of stay, in-hospital mortality, and functional rehabilitation indexes

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# ABSTRACT

*Objective*: To determine the relationships between malnutrition and nutrition-related conditions according to the European Society of Clinical Nutrition and Metabolism (ESPEN) consensus and guidelines and clinical outcomes in postacute rehabilitation.

*Methods*: Of 102 eligible inpatients, 95 (84.5 years old, 63.2% women) fulfilled inclusion criteria: aged  $\geq$ 70 years, body mass index <30 kg/m<sup>2</sup>, admission for rehabilitation. Mini-Nutritional Assessment-Short Form (MNA-SF $\leq$ 11) identified patients "at risk" and ESPEN basic and etiology based definitions were applied. Nutrition-related conditions (sarcopenia, frailty, overweight/obesity, micronutrient abnormalities) were determined. We assessed the relationship between these conditions and the clinical and rehabilitation outcomes (relative functional gain, rehabilitation efficiency) during hospitalization.

*Results*: All patients were "at risk" by MNA-SF criteria and 31 reported unintentional weight loss >5% in the last year or 2–3 kg in the last 6 months. Nineteen fulfilled the ESPEN basic definition, of which 10 had disease-related malnutrition with inflammation and 9 without inflammation, and 20 had cachexia. Sarcopenia (n = 44), frailty (n = 94), overweight/obesity (n = 59), and micronutrient abnormalities (n = 70) were frequent. Unintentional weight loss impaired all functional outcomes and increased length of stay [OR = 6.04 (2.87–9.22); p < 0.001]. In multivariate analysis, relationships between rehabilitation impact indices and the ESPEN basic and etiology-based definitions observed in univariate analysis persisted only (and marginally) for relative functional gain [OR = 13.24 (0.96–181.95); p = 0.005]. Infrequent in-hospital mortality prevented meaningful analysis of this outcome.

*Conclusions:* ESPEN basic and etiology-based definitions and nutrition related disorders were determined in postacute care. Malnutrition was associated with poor rehabilitation outcomes, mainly due to unintentional weight loss.

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Abbreviations: BMI, body mass index; ESPEN, European Society of Clinical Nutrition and Metabolism; EWGSOP, European Working Group on Sarcopenia in Older People; FFMI, fat-free mass index; MNA, Mini-Nutritional Assessment; MNA-SF, Mini-Nutritional Assessment-Short Form; REI, Rehabilitation efficiency index; RFG, relative functional gain; SD, standard deviation; STROBE, Strengthening the Reporting of Observational Studies in Epidemiology; WHO, World Health Organization.

# 1. Introduction

Malnutrition, a common condition in older adults, is associated with adverse medical consequences, contributing to frailty, morbidity, mortality, and use of health resources (Agarwal et al., 2013; Chen et al., 2010; Guyonnet, Secher, & Vellas, 2015; Marshall, 2016; Morley, 2015; Shen et al., 2011; Soysal et al., 2017). Although early identification and management of individuals at risk of malnutrition make it possible to reverse its adverse effects (Cederholm et al., 2015, 2017; Cruz-Jentoft, Kiesswetter, Drey, & Sieber, 2017; Deutz et al., 2014; Deutz et al., 2016; Lindegaard Pedersen, Pedersen, & Damsgaard, 2017; Shen et al., 2011), the condition often remains unrecognized and undermanaged, partly due to the lack of clear and consensual guidelines appropriate for all patients and settings (Cederholm & Jensen, 2017; Cederholm et al., 2015; Hamirudin, Charlton, & Walton, 2016; Lee & Tsai, 2012; Vischer et al., 2012).

The European Society of ClinicalNutrition and Metabolism (ESPEN) has proposed a consensus statement on nutrition for all age ranges and healthcare settings providing a unified, simple and reliable tool for malnutrition diagnosis, independently of etiology (Cederholm et al., 2015; Rojer et al., 2016). This consensus was further completed with the ES-PEN Guidelines on Definition and Diagnoses of Malnutrition, with an etiology-based diagnostic tree including the identification of inflammatory markers, a crucial step for identifying and addressing malnutrition (Cederholm et al., 2017).

The ESPEN consensus has already shown appropriateness for the diagnosis of malnutrition, and engaging in corrective interventions, in a wide range of patients including acutely ill middle-aged patients and geriatric patients hospitalized with diabetes (Rojer et al., 2016; Sanz-Paris et al., 2016) and in postacute care (Sánchez-Rodríguez et al., 2016). However, the relationship of the ESPEN consensus with clinical outcomes during hospitalization in rehabilitation settings is not well established.

The post-acute care pathways are key moments in the patient's trajectory, reflecting their vulnerability, as illustrated by longer hospital stay (Kruizenga et al., 2016), increased inhospital mortality, or poor functional prognosis (Arinzon, Fidelman, Zuta, Peisakh, & Berner, 2005; Cerri et al., 2015; Cruz-Jentoft et al., 2017; Guyonnet et al., 2015; Landi et al., 2002; Nunes, Flores, Mielke, Thumé, & Facchini, 2016; Sánchez-Rodríguez et al., 2014). Thus, since nutritional status is recognized as the most potent prognostic factor in older adults (Sanz-París et al., 2016; Shen et al., 2011), we hypothesize that nutritional disorders according to the ESPEN consensus may be associated with worse post-acute care pathways. Moreover, as the ESPEN basic diagnostic tool is composed of various criteria, including unintentional weight loss, low body mass index (BMI) and low fat-free mass index (FFMI) (Cederholm et al., 2015), an assessment of the respective weight of each criterion and its potential to predict adverse health events seems to be warranted.

The main objective of this study was to determine the relationship, if any, between nutritional disorders diagnosed according to the ESPEN consensus and clinical and functional rehabilitation outcomes; a second objective was to determine if an etiology-based malnutrition diagnosis provides complementary information about the prognosis of older patients during post-acute rehabilitation care.

# 2. Methods

# 2.1. Design

Cross-sectional analysis of consecutive hospitalized older patients who participated in a larger prospective study on sarcopenia and functional outcomes (Sánchez-Rodríguez et al., 2014; Sánchez-Rodríguez, Marco et al., 2015). Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) Guidelines for reporting observational research cohorts were followed (See file 1).

# 2.2. Setting

The study was carried out in a post-acute geriatric rehabilitation care unit in a university hospital in Barcelona (Catalonia, Spain). This post-acute hospitalization unit focuses on rehabilitation and recovery during a defined period of time, usually about two weeks before a scheduled home discharge.

# 2.3. Participants

The study population consisted of 102 consecutive patients who met inclusion criteria during a five-month period: age  $\geq$ 70 years; admitted in the post-acute rehabilitation care unit for functional loss resulting from a non-disabling medical disease. Patients whose general and/or cognitive condition (Mini-Mental Status Examination  $\leq$ 21) prevented completion of the diagnostic tests and an active physical rehabilitation program were excluded.

# 2.4. Procedure

Upon admission to the post-acute rehabilitation unit, patients were screened for risk of malnutrition using the Mini-Nutritional Assessment Short-Form (MNA-SF) (Camina-Martín et al., 2015; Kaiser et al., 2009; Lee & Tsai, 2012). The ESPEN consensus definition of malnutrition was later applied to all screened subjects with MNA-SF score ≤11. According to the ESPEN consensus (Cederholm et al., 2015), the variables that most accurately reflect malnutrition are weight loss, reduced BMI, and reduced FFMI. Unintentional weight loss was determined by patient and caregiver anamnesis and medical records, either by documented weight loss of at least 5% in the previous 12 months during an underlying illness (Evans et al., 2008) and/or by item 11 on the Kihon checklist: "Have you experienced more than 2-3 kg unintentional weight loss over the past 6 months? Yes = 1, No = 0." (Komai et al., 2016). BMI was calculated from weight and height (kg/m<sup>2</sup>). Body weight was measured to the nearest 0.1 kg; height was measured in all patients who were able to stand and a knee height equation was applied in bedridden patients unable to stand safely. Fat-free mass (FFM), expressed in kg, was measured by bioimpedance (Bodystat 1500, Bodystat Ltd., Isle of Man British Isles) as previously described (Sánchez-Rodríguez et al., 2016). The FFM values were divided by height squared to obtain the fat-free mass index (FFMI), expressed in kg/m<sup>2</sup> and compared with those of the reference population (Schutz, Kyle, & Pichard, 2002).

Nutrition-related conditions (sarcopenia, frailty, overweight and obesity, and micronutrient abnormalities) were also considered. Sarcopenia was assessed following European Working Group on Sarcopenia in Older People (EWGSOP) criteria: low muscle mass and low muscle function or low physical performance assessed with bioimpedance analysis, isometric handgrip dynamometry and gait speed in a 4-m walk test as previously described (Cruz-Jentoft et al., 2010; Sánchez-Rodríguez et al., 2014). Gait speed was considered 0 m/s in bedridden patients unable to stand (Cereda, Bertoli, Vanotti, & Battezzati, 2010). Frailty was assessed by the Frailty Phenotype (Fried et al., 2001) when 3 of the following criteria were present: weight loss, weakness, exhaustion, slow walking speed, and low physical activity. Overweight and obesity were considered following World Health Organization recommendations: overweight, BMI 25–30 kg/m<sup>2</sup> and obesity, BMI  $\geq$  30 kg/m<sup>2</sup>. Micronutrient abnormalities: Total proteins, total cholesterol, triglycerides, homocysteine-related markers (folic acid and B12 vitamin), thyroid-stimulating hormone, iron profile (serum ferritin), iron.

ionogram (sodium, potassium), and renal profile (creatinine, urea and glomerular filtration rate from the equation developed by the Modification of Diet in Renal Diseases Study). Following an overnight fast, 10 ml of venous blood were collected under standardized conditions between 7 and 9 am, with the patient in a supine position. Blood handling and collection were carried out under strictly standardized conditions. For purpose of analysis, micronutrient abnormalities were considered as a dichotomous variable to determine whether one or more of the biomarkers of interest was altered.

Patients fulfilling basic diagnostic malnutrition criteria were further considered for the etiology-based diagnoses: Disease-related malnutrition (DRM), with and without inflammation, according to albumin serum levels (reference cut-off point 3.2 mg/dl) (Cederholm et al., 2017) and Malnutrition/undernutrition without disease. Cachexia (wasting disease) diagnostic criteria in adults were applied: weight loss of at least 5% in 12 months or less in the presence of underlying illness, plus three of the following criteria: decreased muscle strength, fatigue (defined as physical and/or mental weariness resulting from exertion), anorexia (total caloric intake <20 kcal/kg body weight/day; <70% of usual food intake), low fat-free mass index, or abnormal biochemistry (hemoglobin <12 g/dl or low serum albumin <3.2 g/dl) (Evans et al., 2008).

Main outcome variables were changes in the functional status assessed with the Rehabilitation Impact Indexes: Relative Functional Gain (RFG, the achieved percentage of potential gain), corrected with premorbid functional status [RFG = (Barthel index at discharge - Barthel index at admission)/(Barthel index premorbid - Barthel index at admission)  $\times$  100]; and the Rehabilitation Efficiency Index (REI), designed to incorporate the duration of therapy [REI = (Barthel index at discharge - Barthel index at admission)/(length of hospital stay)] (Koh, Chen, Petrella, & Thind, 2013; Sánchez-Rodríguez, Miralles et al., 2015). Cut points for these outcomes were RFG <35%, and REI <0.50, respectively. Regarding RFG, for purposes of analysis, negative or indeterminate values resulting from calculations were converted to 0. Patients who died, lost functional capacity during their stay in the unit or were transferred to acute care hospitals for an acute event or worsening clinical status are included in this group (Sánchez-Rodríguez, Miralles et al., 2015). Intrahospital mortality and length of stay (days) were also considered for purpose of analysis.

# 2.5. Ethics

National and international research ethics guidelines were followed, including the Deontological Code of Ethics, Declaration of Helsinki, and Spain's confidentiality law concerning personal data (*Ley Orgánica* 15/1999, 13 December, *Protección de Datos de Carácter Personal*). Detailed, understandable oral and written information was provided to patients and family members, and informed consent to participate was signed by all participants. The study was approved by the local Clinical Ethics Committee (CEIC Parc de Salut Mar, ref. 4447/I).

# 2.6. Statistical analysis

Descriptive analysis of the sample used percentages and frequency distributions for categorical variables, and means (SD) for quantitative continuous variables. Univariate analysis was used to check clinical and functional characteristics of the study participants according to the presence of the basic definition of malnutrition. Qualitative variables were compared by Chi-square or Fisher exact test, as appropriate and quantitative variables by Student *t*-test. Changes in the Barthel index during the study period were assessed by analysis of variance using a repeated-measures mixed design (intrasubject) and a one-factor design (intersubject). As histogram and Q–Q plot showed that length of stay was not normally distributed, median regression was applied to check differences in medians (MD).

For categorical outcomes (RFG, REI), logistic regressions were used to determine Odds Ratios of poorer outcomes. Univariate and multivariate models were performed in order to determine which of the covariates were statistically significant. All covariates included in univariate analysis were also included in the corresponding multivariate model, with malnutrition considered as two distinct variables: disease-related malnutrition with and without inflammation. P values less than 0.05 were considered as statistically significant. Analysis was performed using SPSS 22 (IBM Corporation, SPSS, INC., Chicago, IL, USA).

# 3. Results

Of the 102 patients hospitalized in the rehabilitation unit during the study period, 95 met inclusion criteria (mean age, 84.5 years, standard deviation [SD] 6.5 years; 63.2% women). Two of these patients were excluded from the analysis because they died within the first 48 h of admission, before geriatric assessment was performed. Table 1

#### Table 1

Clinical and functional characteristics of the study participants according to basic malnutrition status (n = 95).

	Total sample ( $n = 95$ )	Malnutrition ( $n = 19$ )	No malnutrition ( $n = 76$ )	р
Age (years)	84.5 (6.5)	84.3 (5.3)	84.6 (6.8)	0.479
Sex (M:W)	35:60	6:13	29:47	0.595
Body mass index (Kg/m <sup>2</sup> )	25.5 (4.3)	21.7 (4.3)	26.3 (3.9)	0.005
Fat-free mass index (FFMI, Kg/m <sup>2</sup> )	14.9 (2.9)	12.7 (1.7)	15.4 (2.9)	0.007
Fat-free mass (Kg)	38.4 (10.3)	32.4 (6.6)	39.7 (10.5)	0.069
Charlson comorbidity index	2.4 (1.8)	2.5 (2.2)	2.3 (1.7)	0.466
Short Portable Mental Status Questionnaire	4.2 (3.1)	5.1 (3.4)	4.0 (3.0)	0.265
Lawton index	2.6 (2.6)	2.5 (2.9)	2.6 (2.5)	0.577
Barthel index:				
-Prior	71.4 (21.6)	66.4 (25.3)	72.5 (20.8)	0.005
-At admission	27.0 (15.4)	19.1 (14.8)	28.7 (15.1)	
-At discharge	54.3 (26.2)	38.9 (29.1)	57.7 (24.5)	
Length of stay (days):				
-In acute care unit	15.0 (12.1)	18.3 (17.0)	15.2 (11.5)	0.805
-In post-acute care unit	14.9 5.8)	18.3 (8.1)	14.1 (4.9)	0.009
Rehabilitation efficiency index	0.4 (0.3)	0.2 (0.2)	0.4 (0.3)	0.008
Relative functional gain (RFG, %)	56.8 (40.7)	33.7 (36.9)	62.7 (39.8)	0.002

Data are expressed as numbers for sex distribution and as mean and standard deviation (SD) for continuous variables.

a Changes in Barthel index during the study period were assessed by analysis of variance using both a repeated-measures mixed design and a one-factor design.

shows the baseline general and geriatric assessment of the participants. Fig. 1 shows the sample distribution according to nutritional disorders and nutrition-related conditions, after applying ESPEN basic diagnoses. In these deconditioned patients, frailty was confirmed in the whole sample by Fried phenotype. Fig. 2 depicts the malnutrition diagnostic tree. All patients were identified as at risk of malnutrition (MNA-SF scores  $\leq$ 11); 19 patients fulfilled the basic criteria for a diagnosis of malnutrition. Etiology-based malnutrition diagnoses completed this tree. Ten of the 19 patients had disease-related malnutrition (DRM) with inflammation due to acute disease and 9 patients had DRM without inflammation. Unintentional weight loss was present in 31 patients. Table 2 shows distribution of weight loss, reduced BMI, and reduced fat-free mass index (FFMI) in patients fulfilling criteria of ESPEN basic diagnosis and malnutrition-related conditions. Reduced FFMI was the most prevalent feature in the total sample and in all the subgroups, including the overweight and obese subgroup of patients.

Clinical characteristics, nutritional disorders according to the ES-PEN consensus (basic diagnoses), nutrition-related conditions, and etiology-based malnutrition diagnosis were further related to poor functional rehabilitation and clinical outcomes, as shown in Table 3 (uni-

variate analysis) and Table 4 (multivariate analysis). Univariate analysis showed that poorer RFG results were associated with unintentional weight loss (OR = 0.16; 95% CI: 0.06–0.42; p < 0.001) and basic diagnosis of malnutrition (OR = 0.12; 95% CI: 0.04-0.37; p < 0.001). The association between RFG and the etiology-based diagnoses was significant for DRM with inflammation (OR = 0.14; 95% CI: 0.03-0.63; p = 0.01) and without inflammation (OR = 0.19; 95% CI: 0.05-0.76; p = 0.019). The same associations were observed in the multivariate analysis for unintentional weight loss (OR = 4.25; 95% CI: 1.08–16.75; p = 0.039) and DRM with inflammation (OR = 13.24; 95% CI: 0.96-181.95; p = 0.05). Finally, a poor REI score was highly associated with unintentional weight loss (OR = 5.06; 95% CI: 1.58–16.19; p = 0.006) and with basic diagnosis of malnutrition (OR = 5.36; 95%) CI: 1.15–24.92; p = 0.032). Only the relationship with unintentional weight loss persisted after multivariate analysis (OR = 4.49; 95% CI: 0.99-20.31; p = 0.051).

Length of stay (Table 5) was associated with unintentional weight loss in univariate (median difference [MD] = 7; 95% CI: 4.29–9.71; p < 0.001) and multivariate analysis (MD = 6.04; 95% CI: 2.87–9.22;

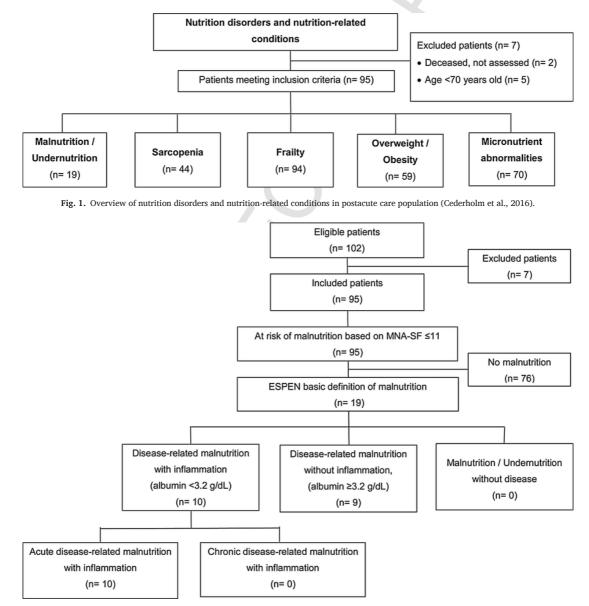


Fig. 2. Diagnostic tree of malnutrition, from identification of risk to basic definition of malnutrition and etiology-based diagnoses (Cederholm et al., 2016).

# Table 2

Unintentional weight loss, reduced body mass index, and reduced fat-free mass index according to malnutrition basic diagnosis and malnutrition-related conditions.(\*) Data are expressed as numbers and percentages.

	Malnutrition (ESPEN) (n = 19)	Cachexia (Evans) (n = 20)	ns)		4	Total sample (n = 95)
			Sarcopenia (EWGSOP) (n = 44)	Frailty (Fried) (n = 94)	Overweight- Obesity (WHO) (n = 59)	
Unintentional weight loss	14 (73.7%)	12 (60%)	13 (29.5%)	31 (33%)	17 (28.8%)	31 (32.6%)
Reduced body mass index (<18.5 kg/m <sup>2</sup> ) Reduced fat-free mass index (<15 kg/m <sup>2</sup> in women, <17 kg/m <sup>2</sup> in men)	4 (21.1%) 19 (100%)	3 (15%) 20 (100%)	3 (6.8%) 43 (97.7%)	4 (4.3%) 59 (62.8%)	0 (0%) 29 (49.2%)	4 (4.2%) 60 (63.2%)

Abbreviations:: European Society of Clinical Nutrition and Metabolism (ESPEN), European Working Group on Sarcopenia in Older People (EWGSOP), World Health Organization (WHO). (\*) Data are expressed as numbers and percentages.

# Table 3

Factors affecting poor rehabilitation outcomes, according to clinical characteristics, components of the basic definition of malnutrition, and etiology-based diagnoses of malnutrition. Univariate analysis.

	Relative functional gain		Rehabilitation efficiency index	
	Odds ratio (95%CI)	p	Odds ratio (95%CI)	р
Clinical characteristics				
Age	1.01 (0.94–1.08)	0.853	1 (0.94–1.07)	0.997
Sex	1.25 (0.49–3.20)	0.642	0.96 (0.39-2.34)	0.923
Comorbidity (Charlson>2)	1.08 (0.83–1.41)	0.559	0.89 (0.70-1.13)	0.339
Unintentional weight loss	0.16 (0.06-0.42)	< 0.001	5.06 (1.58-16.19)	0.006
Albumin < 3.2g/dL	1.02 (0.40-2.60)	0.96	1.11 (0.46–2.68)	0.812
Basic definition of malnutrition and nutrition-related conditions				
Malnutrition	0.12 (0.04-0.37)	< 0.001	5.36 (1.15-24.92)	0.032
Sarcopenia	0.71 (0.28–1.78)	0.465	1.26 (0.53-2.30)	0.603
Overweight-obesity	1.38 (0.55–3.50)	0.494	0.84 (0.34-2.04)	0.694
Micronutrient abnormalities	0.87 (0.34-2.26)	0.783	0.54 (0.22–1.37)	0.198
Etiology-based diagnoses				
Disease-related malnutrition with inflammation	0.14 (0.03–0.63)	0.01	4.364 (0.521-36.57)	0.174
Disease-related malnutrition without inflammation	0.19 (0.05-0.76)	0.019	5 (0.604-41.392)	0.136
Cachexia	0.40 (0.14–1.16)	0.093	1.08 (0.37–3.19)	0.884

# Table 4

Factors affecting poor rehabilitation outcomes, according-clinical characteristics, components of the basic definition of malnutrition, and etiology-based diagnoses of malnutrition. Multivariate analysis.

	Relative functional g	Relative functional gain		Rehabilitation efficiency index	
	Odds ratio (95%CI)	р	Odds ratio (95%CI)	р	
Clinical characteristics					
Age	1.00 (0.92–1.09)	0.987	1.01 (0.93-1.09)	0.859	
Sex	0.64 (0.21–1.99)	0.445	0.78 (0.27-2.27)	0.647	
Comorbidity (Charlson > 2)	0.86 (0.62–1.18)	0.349	0.88 (0.66-1.17)	0.367	
Unintentional weight loss	4.25 (1.08-16.75)	0.039	4.49 (0.99-20.31)	0.05	
Basic definition of malnutrition and nutrition-related conditions					
Malnutrition	_	-	_	-	
Sarcopenia	0.86 (0.49–7.02)	0.362	2.02 (0.62-6.62)	0.245	
Overweight-obesity	1.01 (0.29–3.55)	0.983	0.78 (0.22-2.70)	0.691	
Micronutrient abnormalities	0.81 (0.26-2.50)	0.709	0.46 (0.16–1.34)	0.156	
Etiology-based diagnoses					
Disease-related malnutrition with inflammation	13.24 (0.96–181.95)	0.053	6.82 (0.38-122.08)	0.192	
Disease-related malnutrition without inflammation	5.47 (0.73-40.95)	0.098	4.23(0.29-60.51)	0.289	
Cachexia	0.22 (0.02–2.37)	0.211	0.17 (0.02–1.26)	0.083	

p<0.001). The association between length of stay and DRM without inflammation observed in univariate analysis (MD = 4; 95% CI: 0.39–7.61; p=0.030) was not maintained after multivariate analysis.

Finally, although intrahospital mortality (Table 6) was an unfrequented event (n = 11), it was related to sex in both univariate analy-

sis (OR = 0.07; 95% CI:0.01–0.46; p = 0.006) and multivariate analysis (OR = 0.24; 95% CI: 0.07–0.89; p = 0.032) and with unintentional weight loss in both univariate (OR = 0.003; 95% CI: 0.00–0.31; p = 0.014) and multivariate analysis (OR = 0.16; 95% CI: 0.02–1.28; 0 = 0.084).

## Table 5

Factors affecting length of stay in postacute care, according-clinical characteristics, components of the basic definition of malnutrition, and etiology-based diagnoses of malnutrition. Univariate and multivariate analysis.

	Length of stay in postacute ca	re			
	Univariate analysis		Multivariate analysis		
	MD <sup>a</sup> (95%CI)	р	MD <sup>a</sup> (95%CI)	р	
Clinical characteristics					
Age	-0.06 (-0.22 to 0.10)	0.446	-0.03 (-0.21 to 0.15)	0.747	
Sex	1 (-1.17 to 3.17)	0.362	0.54 (-1.88 to 2.97)	0.656	
Comorbidity (Charlson > 2)	0 (-0.62 to 0.62)	1	0.10 (-0.55 to 0.76)	0.755	
Unintentional weight loss	7 (4.29–9.71)	< 0.001	6.04 (2.87-9.22)	< 0.001	
Basic definition of malnutrition and nutrition-related conditions					
Malnutrition	_	-	_	_	
Sarcopenia	0 (-2.40 to 2.40)	1	0.07 (-2.74 to 2.89)	0.959	
Overweight-obesity	1 (-1.64 to 3.64)	0.454	0.82 (-1.97 to 3.61)	0.559	
Micronutrient abnormalities	0 (-2.77 to 2.77)	1	-0.54 ( $-2.98$ to 1.89)	0.658	
Etiology-based diagnoses					
Disease-related malnutrition with inflammation	0 (-3.77 to 3.77)	1	-2.10 (-7.49 to 3.29)	0.440	
Disease-related malnutrition without inflammation	4 (0.39–7.61)	0.03	4.18 (-0.49 to 8.84)	0.079	
Cachexia	0 (-3.43 to 3.43)	1	-2.34 ( $-6.52$ to 1.85)	0.270	

<sup>a</sup> Median difference.

## Table 6

Factors affecting intrahospital mortality in postacute care, according to clinical characteristics, components of the basic definition of malnutrition, and etiology-based diagnoses of malnutrition. Univariate and multivariate analysis.

	Univariate analysis		Multivariate analysis	
	Odds ratio (95%CI)	Р	Odds ratio (95%CI)	р
Clinical characteristics				
Age	1.11 (0.97–1.27)	0.135	1.1 (0.99–1.22)	0.063
Sex	0.07 (0.01-0.46)	0.006	0.24 (0.07-0.89)	0.032
Comorbidity (Charlson $>2$ )	1.49 (0.91–2.43)	0.108	1.21 (0.88-0.68)	0.238
Unintentional weight loss	0.003 (0.00-0.31)	0.014	0.16 (0.02–1.28)	0.084
Basic definition of malnutrition and nutrition-related conditions				
Malnutrition	-	-	_	-
Sarcopenia	9.19 (0.98–68.11)	0.052	2.69 (0.75–9.64)	0.130
Overweight-obesity	0.69 (0.14–18.86)	0.689	0.85 (0.25-2.91)	0.797
Micronutrient abnormalities	0.59 (0.12–3.0)	0.527	0.85 (0.25-2.91)	0.797
Etiology-based diagnoses				
Disease-related malnutrition with inflammation	23.79 (0.41–1380.89)	0.126	0.84 (0.10-7.39)	0.876
Disease-related malnutrition without inflammation	175.91 (1.80–17153.45)	0.027	0.74 (0.08-6.40)	0.782
Cachexia	0.767 (0.04–16.58)	0.87	1.27 (0.31-5.23)	0.736

# 4. Discussion

Our study applied both the ESPEN consensus and guidelines to describe the influence of basic and etiology-based diagnoses of malnutrition on clinical outcomes during hospitalization for postacute care. As malnutrition and disability are linked bidirectionally to mobility and disability (Landi et al., 2017), an early identification of malnourished patients would help the multidisciplinary care team to set realistic aims, plan therapeutic strategies, and provide the patient and caregivers with more precise information. The influence of the ESPEN consensus on clinical outcomes was recently reported in geriatric patients with diabetes in acute care (Sanz-París et al., 2016), but its influence on clinical and rehabilitation outcomes during hospitalization in postacute care has not previously been described

In our study, malnutrition and nutritional disorders are associated with worse functional prognosis in older patients during postacute rehabilitation care. This association is probably explained by weight loss prior to admission, which has been the most powerful predictor of poor functional outcomes in postacute care in our study and in previous findings (Lee & Tsai, 2012; Morley, 2015). Unintentional weight loss has been proposed as a key indicator to assess formal nutrition because of its validity, feasibility, efficiency, and availability for every population and level of healthcare assistance (Cederholm & Jensen, 2017). Nevertheless, our results highlight that reduced FFMI is the most powerful indicator of malnutrition, sarcopenia, frailty, and cachexia.

Although the influence of BMI on mortality is still a controversial issue, most authors agree that a protective optimal BMI range of 25–30 kg/m<sup>2</sup> provides better nutritional status and protects older individuals from diseases and the malnutrition caused by diseases (Chang, Beason, Hunleth, & Colditz, 2012). In our sample, overweight and obese patients were analyzed as one group because no significant differences in age, functional and cognitive status, comorbidity, length of stay, rehabilitation impact indexes, or survival were observed between them.

Our study also evaluates micronutrient abnormalities and albumin serum levels, and sheds light on the updated role of these biomarkers, which have been commonly used as markers of malnutrition in clinical settings (Cabrerizo et al., 2015) even though their relation with malnutrition was controversial (Reijnierse et al., 2015). Evidence-based medicine agrees on considering serum albumin and C-reactive protein as markers of inflammation (Bano et al., 2017; Cederholm et al., 2017; Reijnierse et al., 2015; Schaap et al., 2009), but not as useful tools for basic diagnosis of malnutrition. Nevertheless, they have become a part of the etiology-based diagnoses of malnutrition since the ESPEN diagnostic tree was proposed as the gold standard for nutritional assessment in Europe.

Although unintentional weight loss and malnutrition with and without inflammation was associated with the rehabilitation outcomes in the univariate analysis, only unintentional weight loss emerged as a significant risk factor of poor rehabilitation outcomes, longer hospital stay, and 2-year mortality. In a large multicenter study in a cohort of older diabetic patients hospitalized for acute care (Sanz-París et al., 2016), malnutrition as defined by the ESPEN consensus statement was the only factor associated within hospital mortality. In our postacute care sample, the impact of malnutrition on mortality could not be determined because mortality was a rare event. Further research is needed on this issue in larger samples or in a follow-up after hospital discharge.

Our results support calls for simplified diagnostic tools that connect clinical and basic sciences and act bidirectionally, summarized as the 'Action Research' philosophy: from bedside to bench to bedside again, providing both meaningful challenges and new answers for clinicians and researchers (Beauchet, Fantino, & Annweiler, 2012). The findings in our postacute care unit may contribute to efforts to reach an international unified definition of malnutrition for application in all geriatric settings (Cederholm & Jensen, 2017).

# 4.1. Limitations of the study

There was an initial selection bias, given that patient selection was based on the potential to pursue rehabilitation therapy. Those who were unable to walk or needed help from two or more persons prior to their hospitalization, as well as patients who do not require rehabilitation after the acute medical process, are not admitted to the post-acute care unit. The relatively small number of patients is also a potential limitation. The use of the MNA-SF as a screening tool is valid (Kaiser et al., 2009; Lee & Tsai, 2012) and has been recommended by the Spanish Geriatrics and Gerontology Society (Camina-Martín et al., 2015), but using the full MNA might have improved specificity. Other potential limitations of the study are related to the REI; this index is highly conditioned by the mean length of stay, which could be influenced by multiple factors that do not depend on the rehabilitation process (Sánchez-Rodríguez, Miralles et al., 2015). Anthropometric measurements, such as the use of demispan or knee height in patients unable to stand (12 patients) without taking into account possible kyphosis or vertebral osteoporotic degenerative changes, might interfere with an accurate BMI, FFMI, basic diagnosis of malnutrition, and sarcopenia or cachexia diagnosis (Bahat et al., 2012; Sánchez-Rodríguez et al., 2016). Limitations for determining weight loss in older populations have been well described (Robbins, 1989), and could be avoided by systematic measurement and recording as part of the comprehensive geriatric assessment. Regarding the etiology-based diagnoses, only albumin levels were considered as markers of inflammation, when C-reactive protein is likely a more specific marker. Last, but not least, the observational design of the study might be considered a limitation; further studies should be designed to assess causality and associations related with postdischarge and follow-up events.

# 4.2. Conclusions and future lines of clinical research

The key point of this study is to discuss practical implications of malnutrition assessment. Further research is urgently needed to assess the value of a basic diagnosis of malnutrition, according to the ESPEN consensus statement and its subscores, as a predictor of adverse outcomes after hospital discharge, including institutionalization, unexpected hospital readmissions, and mortality.

# **Conflict of interest**

None.

## Author contributions

DS, EM conceived and designed the experiments; DS, EM, NR, AT performed the experiments; DS, EM, XD, CA analyzed and interpreted the data; DS, EM, CA, NR, OVI, FE, JMM contributed reagents, materials, analysis tools or data; DS, EM, CA wrote the paper.

# Uncited reference

Landi et al. (2015).

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