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ESPEN Endorsed Recommendation

The GLIM criteria for the diagnosis of malnutrition — A consensus report from the global clinical nutrition community

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SUMMARY

Rationale: This initiative is focused on building a global consensus around core diagnostic criteria for malnutrition in adults in clinical settings.

Methods: In January 2016, the Global Leadership Initiative on Malnutrition (GLIM) was convened by several of the major global clinical nutrition societies. GLIM appointed a core leadership committee and a supporting working group with representatives bringing additional global diversity and expertise. Empirical consensus was reached through a series of face-to-face meetings, telephone conferences, and e-mail communications.

Results: A two-step approach for the malnutrition diagnosis was selected, i.e., first screening to identify "at risk" status by the use of any validated screening tool, and second, assessment for diagnosis and grading the severity of malnutrition. The malnutrition criteria for consideration were retrieved from existing approaches for screening and assessment. Potential criteria were subjected to a ballot among the GLIM core and supporting working group members. The top five ranked criteria included three phenotypic criteria (non-volitional weight loss, low body mass index, and reduced muscle mass) and two etiologic criteria (reduced food intake or assimilation, and inflammation or disease burden). To diagnose malnutrition at least one phenotypic criterion and one etiologic criterion should be present. Phenotypic metrics for grading severity as Stage 1 (moderate) and Stage 2 (severe) malnutrition are proposed. It is recommended that the etiologic criteria be used to guide intervention and anticipated outcomes. The recommended approach supports classification of malnutrition into four etiology-related diagnosis categories. Conclusion: A consensus scheme for diagnosing malnutrition in adults in clinical settings on a global scale is proposed. Next steps are to secure further collaboration and endorsements from leading nutrition professional societies, to identify overlaps with syndromes like cachexia and sarcopenia, and to promote dissemination, validation studies, and feedback. The diagnostic construct should be re-considered every 3-5 years. © 2018 Elsevier Ltd, European Society for Clinical Nutrition and Metabolism and American Society for Parenteral and Enteral Nutrition. All rights reserved.

1. Introduction

Malnutrition due to disease, poverty, hunger, war, and natural catastrophe is a fate suffered by greater than 1 billion of the world's population. Historically, starvation and famine were prevalent causes of malnutrition and they remain so today. However, with improvements in agriculture, education, public health, healthcare, and living standards, nutrition disorders and related conditions now encompass the full scope of undernutrition, micronutrient abnormalities, obesity, cachexia, sarcopenia, and frailty [1,2].

Malnutrition, e.g. undernutrition, may be caused by compromised intake or assimilation of nutrients but there is growing appreciation that malnutrition may also be caused by disease-associated inflammatory or other mechanisms. The malnutrition that is associated with disease or injury invariably consists of a combination of reduced food intake or assimilation and varying degrees of acute or chronic inflammation, leading to altered body composition and diminished biological function [1–3]. Inflammation contributes to malnutrition through associated anorexia and decreased food intake as well as altered metabolism with elevation of resting energy expenditure and increased muscle catabolism. Altered body composition manifests as a decrease in any marker of muscle mass (fat-free mass, muscle mass index or body cell mass). Thus, malnutrition is associated with adverse functional and clinical outcomes.

Although malnutrition is a global concern associated with incremental morbidity, mortality, and cost, there has been a fundamental lack of consensus on diagnostic criteria for application in clinical settings. No single existing approach has secured broad global acceptance [1,4–8]. Our evolving understanding of the contributions of disease/inflammation may render some concepts of malnutrition in the current International Classifications of Diseases (ICD-10) (http://www.who.int/classifications/icd/en/)

inconsistent with approaches or nomenclature that are currently used in clinical practice and research. Thus, there is an urgent need to establish a global consensus to be used in clinical care settings for adults.

In order to respond to the needs of the clinical nutrition and medical communities the Global Leadership Initiative on Malnutrition (GLIM) was convened in January 2016. GLIM has engaged several of the clinical nutrition societies with global reach to focus on standardizing the clinical practice of malnutrition diagnosis. We also sought to clarify overlaps with related disease classifications including cachexia. The purpose of this specific initiative is to reach global consensus on the identification and endorsement of criteria for the diagnosis of malnutrition in clinical settings.

2. Methods

2.1. The consensus procedure

On January 19, 2016 the Global Leadership Conversation: Addressing Malnutrition was held at the ASPEN Conference [9]. Key breakthroughs at that meeting led to the development of GLIM:

- 1. It was recognized that there was considerable consensus among stakeholders around many malnutrition diagnosis issues
- 2. There was strong commitment for reaching broader global consensus in defining and characterizing malnutrition
- 3. A core leadership committee with representatives of several of the global clinical nutrition societies; ASPEN (www.nutritioncare.org), ESPEN (www.espen.org), FELANPE (www.felanpeweb.org) and PENSA (www.pensa-online.org) was constituted to form GLIM. The core GLIM leadership committee then created a larger supporting working group comprised of

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invited members that brought additional global diversity and expertise to the consensus effort.

4. It was agreed that a series of face-to-face meetings, telephone conferences, and email communications would be used to delineate the GLIM approach.

The first full meeting of the GLIM extended working group was held September 19, 2016 at the ESPEN Congress [10]. Highlighted objectives included consensus development of evidence-based criteria suitable to diverse clinical settings, global dissemination of consensus criteria, and the priority to seek adoption by leading diagnosis classification and coding entities across the globe. It was also agreed that the desired approach to malnutrition diagnosis should be simple and include clinically relevant diagnostic criteria that will be appropriate for application by all healthcare professionals using methods that are widely available. The intent was also to promote global use of consensus criteria that can be readily used with other approaches and additional criteria of regional preference.

3. Results

Consensus was gradually achieved over the course of the GLIM meetings held February 20, 2017 at the ASPEN Conference [11], September 11, 2017 at the ESPEN Congress, and January 25, 2018 at the ASPEN Conference. Meanwhile, discussions were also held with the leadership of The Society of Sarcopenia, Cachexia and Wasting Disorders (SCWD).

3.1. A two-step model for risk screening and diagnosis assessment

There was strong consensus that the key first step in the evaluation of nutritional status is malnutrition risk screening to identify "at risk" status by the use of any validated screening tool [12-14]; some of these tools are noted in Table 1 and the Appendix. This is followed by the second step of assessment for diagnosis and severity grading as described below.

3.2. Criteria selected for malnutrition diagnosis

A comprehensive survey of existing approaches used in screening and assessment of malnutrition was conducted to identify criteria worthy of consideration (Table 1 and the Appendix). It was recognized that these approaches incorporate multiple common criteria. For example, the presence of weight loss and disease burden or inflammation is common to most of them as is reduced food intake (Table 1). Potential consensus criteria from existing approaches as well as additional criteria suggested by participants were subject to further consideration.

In order to establish consensus and endorsement of a minimum set of diagnostic criteria by the core leadership committee and the supporting working group a formal ballot was administered whereby participants ranked proposed diagnosis criteria. The top 5 ranked criteria by an overwhelming majority of GLIM participants were as follows:

- Non-volitional weight loss
- Low body mass index (BMI)
- Reduced muscle mass
- Reduced food intake or assimilation
- Disease burden/inflammation

3.3. Non-volitional weight loss

There was strong GLIM consensus for the inclusion of nonvolitional weight loss as a phenotypic criterion. Validity is well established and there is a robust literature on which threshold selection could be based (Appendix). There must be priority to obtain repeated weight measures over time to identify trajectories of decline, maintenance, and improvement. GLIM participants felt that it is especially important to recognize the pace of weight loss early in the course of disease or injury and to highlight that many patients will have lost appreciable weight prior to presenting to the healthcare.

3.4. Low BMI

There is substantial regional variation in the use of low BMI as a phenotypic criterion for malnutrition diagnosis. North American GLIM representatives indicated that low BMI is seldom used as a clinical malnutrition marker in those regions. The experience from the current American population is that people are often overweight or obese and would need to lose substantial weight before low BMI designation would occur. Since other regions of the world currently make use of BMI as a criterion for recognition of malnutrition, the GLIM consensus includes low BMI. Further research is however needed to secure consensus reference BMI data for Asian populations in clinical settings.

3.5. Reduced muscle mass

Reduced muscle mass is a phenotypic criterion with strong evidence to support its inclusion in the GLIM consensus criteria. However, there is not consensus regarding how best to measure and define reduced muscle mass, particularly in clinical settings. Therefore, GLIM recommends measurement by dual-energy absorptiometry or other validated body composition measures such as bioelectrical impedance, ultrasound, computed tomography or magnetic resonance imaging, but these methods are still not available in most settings for nutritional assessment throughout the globe. Physical examination or anthropometric measures of calf or arm muscle circumference are therefore included as alternative measures. Recommendations are likely to evolve as portable and less costly body composition technologies are developed and become widely available.

For the purpose of recommended cut-off values for muscle mass reductions, GLIM refers to recommendations from the European Working Group on Sarcopenia in Older People (EWGSOP) [15] and from The Foundation of National Institute of Health (FNIH) initiative [16], and the Asian Working Group on Sarcopenia (AWGS) [17]. Reference standards for muscle mass may warrant adjustment for race and sex. Additional research is warranted to establish general reference standards as well as for some specific populations, e.g. in Asia. Examples of recommended thresholds are found in Table 2.

Assessment of muscle function using grip strength or other validated procedures is recommended as a supportive measure in the GLIM consensus (Tables 3 and 4). Decline in muscle strength generally exceeds changes in muscle size [18]. However, irrespective of etiology, appreciable loss of muscle mass is generally accompanied by reduced muscle function. In situations where muscle mass cannot be readily assessed then muscle strength, e.g. hand grip strength, is an appropriate supporting proxy.

3.6. Reduced food intake or assimilation

Reduced food intake is a well-established etiologic criterion for malnutrition that has strong validity. It can have multiple causes including poor oral health, medication side effects, depression, dysphagia, gastrointestinal complaints, anorexia and inadequate

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Table 1 Survey of existing approaches used in screening and assessment of malnutrition and cachexia.

	NRS-2002 [12] ^a	MNA-SF [21] ^{a,b}	MUST [22] ^a	ESPEN 2015 [8] ^a	ASPEN/AND [7] ^a	SGA [4] ^a	Evans 2008 [5] ^c	PEW 2008 [23] ^d	Fearon 2011 [6] ^c
Etiologies									
Reduced food intake	X	X	X	X	X	X		X	X
Disease burden/inflammation	X	X	X	X	X	X	X	X	X
Symptoms									
Anorexia		X				X	X		X
Weakness		X				X	X		
Signs/Phenotype									
Weight loss	X	X	X	X	X	X	X	X	X
Body mass index	X	X	X	X			X	X	X
Lean/fat free/muscle mass		X		X	X	X	X	X	X
Fat mass					X	X		X	
Fluid retention/ascites					X	X			
Muscle function; e.g. grip strength					X	X	X		
Biochemistry							X	X	

NRS-2002: Nutritional Risk Screening-2002, MNA-SF = Mini Nutritional Assessment-Short Form, MUST = Malnutrition Universal Screening Tool, ESPEN = European Society for Clinical Nutrition and Metabolism, ASPEN = American Society of Parenteral and Enteral Nutrition, AND = Academy of Nutritiona and Dietetics, SGA=Subjective Global Assessment, PEW=Protein Energy Wasting.

- Malnutrition approach.
- b Adapted for older adults.
- Cachexia approach.
- d Adapted for chronic kidney disease.

Table 2 Examples of recommended thresholds for reduced muscle mass.

	Males	Females
Appendicular Skeletal Muscle Index (ASMI, kg/m²) [20] Appendicular Skeletal Muscle Index (ASMI, kg/m²) [17] a	<7.26	<5.25
DXA	<7	<5.7
BIA	<7	< 5.4
Fat free mass index (FFMI, kg/m ²) [8]	<17	<15
Appendicular lean mass (ALM, kg) [21] Appendicular lean mass adjusted for BMI = ALM/BMI [21]	<21.4 <0.725	<14.1 <0.591

DXA = dual energy x-ray absorptiometry, BIA = bioelectrical impedance analysis. BMI = body mass index.

nutrition support. Thresholds for relevant impairment of food intake are widely reported (Appendix) and GLIM participants sought to empirically provide a practical synthesis. Reduced assimilation of food/nutrients is associated with malabsorptive disorders like short bowel syndrome, pancreatic insufficiency and after bariatric surgery. It is also associated with disorders like esophageal strictures, gastroparesis, and intestinal pseudoobstruction, as well as with gastrointestinal symptoms like dysphagia, nausea, vomiting, diarrhea, constipation, and abdominal pain. These symptoms have been incorporated as supportive indicators into this GLIM consensus criterion to help to identify poor food intake or assimilation.

Phenotypic and etiologic criteria for the diagnosis of malnutrition.

Phenotypic Criteria ^g		Etiologic Criteria ^g		
Weight loss (%)	Low body mass index (kg/m ²)	Reduced muscle mass ^a	Reduced food intake or assimilation ^{b,c}	Inflammation ^{d-f}
>5% within past 6 months, or >10% beyond 6 months	<pre><20 if < 70 years, or <22 if >70 years Asia: <18.5 if < 70 years, or <20 if >70 years</pre>	Reduced by validated body composition measuring techniques ^a	\leq 50% of ER > 1 week, or any reduction for >2 weeks, or any chronic GI condition that adversely impacts food assimilation or absorption ^{0,c}	Acute disease/injury ^{d,f} or chronic disease-related ^{e,f}

GI = gastro-intestinal, ER = energy requirements.

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^a Recommendations from Asian Working Group for Sarcopenia (AWGS) for Asians.

^a For example fat free mass index (FFMI, kg/m²)) by dual-energy absorptiometry (DXA) or corresponding standards using other body composition methods like bioelectrical impedance analysis (BIA), CT or MRI. When not available or by regional preference, physical examination or standard anthropometric measures like mid-arm muscle or calf circumferences may be used. Thresholds for reduced muscle mass need to be adapted to race (Asia). Functional assessments like hand-grip strength may be considered as a supportive measure.

Consider gastrointestinal symptoms as supportive indicators that can impair food intake or absorption e.g. dysphagia, nausea, vomiting, diarrhea, constipation or abdominal pain. Use clinical judgment to discern severity based upon the degree to which intake or absorption are impaired. Symptom intensity, frequency, and duration should be noted.

Reduced assimilation of food/nutrients is associated with malabsorptive disorders like short bowel syndrome, pancreatic insufficiency and after bariatric surgery. It is also associated with disorders like esophageal strictures, gastroparesis, and intestinal pseudo-obstruction. Malabsorption is a clinical diagnosis manifest as chronic diarrhea or steatorrhea. Malabsorption in those with ostomies is evidenced by elevated volumes of output. Use clinical judgment or additional evaluation to discern severity based upon frequency, duration, and quantitation of fecal fat and/or volume of losses.

Acute disease/injury-related. Severe inflammation is likely to be associated with major infection, burns, trauma or closed head injury. Other acute disease/injury-related conditions are likely to be associated with mild to moderate inflammation.

Chronic disease-related. Severe inflammation is not generally associated with chronic disease conditions. Chronic or recurrent mild to moderate inflammation is likely to be associated with malignant disease, chronic obstructive pulmonary disease, congestive heart failure, chronic renal disease or any disease with chronic or recurrent Inflammation. Note that transient inflammation of a mild degree does not meet the threshold for this etiologic criterion.

C-reactive protein may be used as a supportive laboratory measure.

g Requires at least 1 phenotypic criterion and 1 etiologic criterion for diagnosis of malnutrition.

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Thresholds for severity grading of malnutrition into Stage 1 (Moderate) and Stage 2 (Severe) malnutrition.

	Phenotypic Criteria ^a			
	Weight loss (%)	Low body mass index (kg/m²) ^b	Reduced muscle mass ^c	
Stage 1/Moderate Malnutrition (Requires 1 phenotypic criterion that meets this grade) Stage 2/Severe Malnutrition (Requires 1 phenotypic criterion that meets this grade)	5–10% within the past 6 mo, or 10–20% beyond 6 mo >10% within the past 6 mo, or >20% beyond 6 mo	<20 if <70 yr, <22 if ≥70 yr <18.5 if <70 yr, <20 if ≥70 yr	Mild to moderate deficit (per validated assessment methods – see below) Severe deficit (per validated assessment methods – see below)	

a Severity grading is based upon the noted phenotypic criteria while the etiologic criteria described in the text and Fig. 1 are used to provide the context to guide inter-

3.7. Disease burden/inflammation

GLIM members recognized that disease burden/inflammation has become a widely accepted etiologic criterion in existing screening and assessment tools (Table 1). Clinical diagnosis provides a simple approach to recognition of severe, chronic or frequently recurrent inflammation [1,2,19]. For example, major infections, burns, trauma, and closed head injury are associated with acute inflammation of a severe degree. Indicators of inflammation may include fever, negative nitrogen balance, and elevated resting energy expenditure. Most chronic organ diseases, like congestive heart failure, chronic obstructive pulmonary disease, rheumatoid arthritis, chronic kidney or liver disease and cancer, are associated with chronic or recurrent inflammation of a mild to moderate degree. While severe inflammation is generally easy to discern, clinical judgment is often required to recognize that of lesser degree. Supportive proxy measures of inflammation can include laboratory indicators like serum C-reactive protein (CRP), albumin, or pre-albumin.

3.8. Approach to using combined phenotypic and etiologic criteria for malnutrition diagnosis

Weight loss, reduced BMI, and reduced muscle mass were categorized as phenotypic criteria, and reduced food intake/ assimilation and disease burden/inflammation as etiologic criteria (Table 3 and Fig. 1). For the diagnosis of malnutrition, GLIM recommends that the combination of at least one phenotypic criterion and one etiologic criterion is required (Table 3 and Fig. 1). The selection of threshold values for the consensus diagnostic criteria was guided by review of existing approaches used in screening and assessment as was the selection of threshold values for severity grading described below (see Appendix). The selected threshold values for diagnosis of malnutrition are shown in Table 3. While only the phenotypic criteria are proposed for the severity grading that follows, the inclusion of the etiologic criteria for malnutrition diagnosis is deemed a priority to guide appropriate intervention and anticipated outcomes.

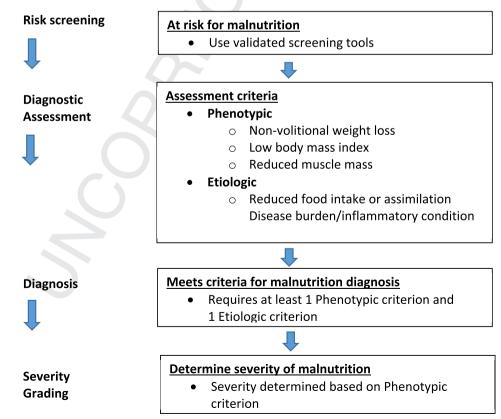


Fig. 1. GLIM diagnostic scheme for screening, assessment, diagnosis and grading of malnutrition.

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Further research is needed to secure consensus reference BMI data for Asian populations in clinical settings.

^c For example appendicular lean mass index (ALMI, kg/m²) by dual-energy absorptiometry or corresponding standards using other body composition methods like bioelectrical impedance analysis (BIA). CT or MRI, When not available or by regional preference, physical examination or standard anthropometric measures like mid-arm muscle or calf circumferences may be used. Functional assessments like hand-grip strength may be used as a supportive measure [15].

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3.9. Severity grading of malnutrition

It is clinically useful to categorize the severity of malnutrition depending on the degree of aberration from established thresholds. Suggested phenotypic metrics for grading severity as Stage 1 (moderate) and Stage 2 (severe) malnutrition are shown in Table 4.

3.10. Etiology-based diagnosis classification

An etiology-based diagnosis classification is endorsed by GLIM consistent with those suggested previously by the International Consensus Guideline Committee [1], the AND/ASPEN Guidelines [7], and the ESPEN Guidelines [2]. The classification includes malnutrition related to chronic disease with inflammation, malnutrition related to chronic disease with minimal or no perceived inflammation, malnutrition related to acute disease or injury with severe inflammation, and malnutrition related to starvation including hunger/food shortage associated with socioeconomic or environmental factors (Table 5).

4. Discussion

This GLIM initiative targets the priority to adopt global consensus criteria so that malnutrition prevalence, interventions, and outcomes may be compared throughout the world. A common malnutrition "language" is a paramount necessity in order to support the development of global standards of care that will promote improved outcomes. The proposed approach for diagnosing malnutrition is based upon a strong consensus endorsing core phenotypic and etiologic criteria that are already in widespread use throughout the world. The intent is to promote global use of these criteria that may in turn be readily used with other approaches and additional criteria of regional preference. The consensus criteria are intended to be simple and readily applied by clinicians and other health practitioners using tools and methods that are readily available. Only modest training should be required. The proposed approach encompasses risk screening and diagnosis but does not entail the robust detail of comprehensive nutrition assessment. It will provide a malnutrition diagnosis that may then be complemented by more comprehensive assessments to provide the basis for individualized care and treatment plans. Consultation of skilled nutrition practitioners like dietitians is recommended for comprehensive assessment based upon regional preferences and availability. Repeated criterion measures over time are recommended so that trajectories of decline, maintenance, and improvement may be identified.

The recommended GLIM approach encompasses both phenotypic and etiologic criteria for the diagnosis of malnutrition but uses only phenotypic criteria cut-points to provide for severity grading. While etiology has not generally been included in criteria supporting the diagnosis of medical conditions in the ICD construct, the inclusion of etiology has been widely adopted in the clinical nutrition community because it serves to guide appropriate interventions and expected outcomes [1]. For example, the presence of disease-associated inflammatory response has potential for major impacts upon treatment approach and anticipated outcome. The GLIM approach acknowledges the diversity and the multi-factorial etiologies

Table 5 Diagnosis category according to underlying etiology.

Malnutrition related to

- Chronic disease with inflammation
- Chronic disease with minimal or no perceived inflammation
- · Acute disease or injury with severe inflammation
- Starvation including hunger/food shortage associated with socio-economic or environmental factors

underlying the development of the malnourished phenotype irrespective of body morphology - lean, normal or obese.

Impairment of muscle strength and function are core phenomena in conditions like sarcopenia [15,16], cachexia [5,6], and frailty [20]. Assessment of muscle strength should be an integral measure in assessment of patients with suspected sarcopenia since impairment of muscle strength is now recognized as a key component for diagnosis of sarcopenia [15,16]. Though inflammatory mediators and other mechanisms besides malnutrition are at play, it is recommended that the GLIM consensus criteria be applied to diagnose malnutrition in persons with sarcopenia, cachexia, and frailty so that the priority to undertake appropriate nutrition interventions may be recognized. The most helpful approaches for these conditions will however require combined multimodal interventions beyond nutritional supplements, like pharmacological agents and exercise.

Similarly, patients with cachexia will meet GLIM consensus criteria for malnutrition related to chronic disease with inflammation. Since there is concern that inclusion of cachexia with other disease-related malnutrition conditions may diminish appreciation for some distinctive features of cachexia, there has been understandable hesitation by some to equate cachexia with this GLIM diagnosis category. The GLIM consensus criteria for malnutrition are therefore intended to be used in parallel with established concepts and nomenclature, including for example, those of cachexia, sarcopenia and frailty.

5. Conclusion

A strong GLIM consensus endorsed the selected core phenotypic and etiologic criteria that are already in widespread use throughout the world. Many studies provide clear evidence that the agreed upon criteria for diagnosis of malnutrition are highly relevant and each of them alone is able to predict adverse clinical outcomes. Since these criteria may be readily used with other approaches and additional criteria of regional preference, their global adoption is more likely. As the initiative moves forward the creation of databases that use the selected criteria will facilitate the comparison of malnutrition prevalence, interventions, and outcomes throughout the world. Such observations can be used to support the development of global standards of care that will promote improved outcomes.

After the launch of the GLIM consensus it is important that the nutrition community use the criteria both in prospective and retrospective cohort studies as well as clinical trials in order to validate its relevance for clinical practice. Next steps are to secure endorsements from leading nutrition professional societies and to promote dissemination, validation testing, and feedback. The GLIM consensus should be re-evaluated based upon review of new studies and advances in screening and assessment every 3-5 years. We will also seek to share the GLIM consensus recommendations with the World Health Organization in the context of the International Classification of Diseases revision process (ICD-11). This is a high priority since this classification scheme guides clinical diagnosis and reimbursement across much of the world. The proposed GLIM consensus criteria target adults in clinical settings but it will also be a priority to work with the World Health Organization and the United Nations to explore the potential for use in other global settings like famine.

Conflict of interest

None declared.

Uncited reference

[24], [25].

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Appendix

Appendix Table 1Cut-offs suggested in the major screening tools.

	Phenotypic criteria			Etiologic criteria		
	Weight loss	Low body mass index (kg/m²)	Reduced muscle mass/muscle function	Reduced food intake	Severe disease/inflammation	
NRS-2002 [12]						
Mild	>5% in 3 mo	NS	NA	50-75% of required preceding week	E.g. hip fracture, chronic disease	
Moderate	>5% in 2 mo	18.5-20.5	NA	25-60% of required preceding week	E.g. major abdominal surgery, stroke	
Severe	>5% in 1 mo	<18.5	NA	0–25% of required preceding week	E.g. head injury, bone marrow transplantation, intensive care	
MNA-SF ^a [21]					,	
Mild	1-3 kg in last months	21-23	NS	NS	NS	
Moderate	"Does not know"	19-21	"Does not go out"	Moderate loss of appetite past 3 mo	Mild dementia	
Severe	>3 kg last months	<19	Bed or chair bound	Severe loss of appetite past 3 mo	Acute disease past 3 mo, or severe dementia/depression	
MUST [22]						
Medium risk	5-10% in 3-6 mo	18.5-20	NA	NS	NA	
High risk	>10% in 3-6 mo	<18.5	NA	Acute illness AND no food intake for >5 d	NA	

NRS-2002 = Nutritional Risk Screening-2002, MNA-SF = Mini Nutritional Assessment-Short Form, MUST = Malnutrition Universal Screening Tool, NA = not applicable, NS = not specified.

Appendix Table 2Cut-offs suggested in major diagnostic tools for malnutrition and cachexi

	Phenotypic criteria			Etiologic criteria		
	Weight loss (%)	Low body Mass Index (kg/m²)	Reduced muscle mass/muscle function	Reduced food intake	Severe disease/inflammation	
SGA [4]						
Moderate/Stage B	5-10% past 6 mo	NA	Mild to moderate deficits in function or muscle mass	"Definite decrease"	Yes	
Severe/Stage C	>10% past 6 mo	NA	Severe deficit in function and muscle mass	"Severe deficit"	Yes	
Evans 2008 [5]						
Cachexia	>5% in <12 mo	<20	Low FFMI, decreased muscle strength	"Anorexia"	Increased CRP/IL6, low serum albumin (<3.2 g/l)	
PEW 2008 [23]						
Protein-energy wasting	>5% in 3 mo, or >10% in >6 mo	<23	Muscle mass down by 5% last 3 mo, or >10% in >6 mo. Reduced MAC	Energy intake <25 kcal/kg BW/d for >2 mo	Chronic kidney disease, Serum albumin <3.8 g/dl	
Fearon 2011 [6]						
Precachexia	<5%	NA	NA	"Anorexia"	Metabolic change	
Cachexia	>5% in 6 mo (>2%)	<20 (when WL>2%)	Sarcopenia - ASMI 7.26/5.45 kg/m 2 (m/w) when WL >2%	"Often reduced food intake"	Cancer with catabolic drive (systemic inflammation)	
ASPEN/AND 2012 [7					,	
Moderate	1–2% in 1 w to 20% in 1 y	NA	Mild muscle loss	<75% of ER for 7 d-3 mo	Yes	
Severe	>2% in 1 week to >20% in 1 year	NA	Moderate to severe muscle loss, or reduced grip strength	<50% of ER for >5 d-<1 mo	Yes	
ESPEN 2015 [8]						
Malnutrition	>5% past 3 mo, or >10%	<18.5, or <20 (<70 y)/ <22 (>70 y)	FFMI <15 kg/m ² (f), <17 kg/m ² (m)	According to any validated tool	NA	

 $SGA=Subjective\ Global\ Assessment,\ NA=not\ applicable,\ NS=not\ specified,\ WL=weight\ loss,\ PEW=protein\ energy\ wasting,\ MAC=mid-arm\ circumference,\ ASMI=appendicular\ skeletal\ muscle\ index\ from\ DEXA,\ FFMI=fat\ free\ mass\ index.$

References

- [1] Jensen GL, Mirtallo J, Compher C, Dhaliwal R, Forbes A, Grijalba RF, et al., International Consensus Guideline Committee. Adult starvation and disease-related malnutrition: a proposal for etiology-based diagnosis in the clinical
- practice setting from the International Consensus Guideline Committee. J Parenter Enter Nutr 2010;34:156–9. Clin Nutr 2010;29:151-3.
- [2] Cederholm T, Barazzoni R, Austin P, Ballmer P, Biolo G, Bischoff SC, et al. ESPEN guidelines on definitions and terminology of clinical nutrition. Clin Nutr 2017;36:49–64.

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^a Adapted for older adults (>65 y).

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- [3] Soeters PB, Reijven PL, van Bokhorst-de van der Schueren MA, Schols JM, Halfens RJ, Meijers JM, et al. A rational approach to nutritional assessment. Clin Nutr 2008;27:706–16.
- [4] Detsky AS, McLaughlin JR, Baker JP, Johnston N, Whittaker S, Mendelson RA, et al. What is subjective global assessment of nutritional status? J Parenter Enter Nutr 1987;11:8–13.
- [5] Evans WJ, Morley JE, Argilés J, Bales C, Baracos V, Guttridge D, et al. Cachexia: a new definition. Clin Nutr 2008;27:793–9.
- [6] Fearon K, Strasser F, Anker SD, Bosaeus I, Bruera E, Fainsinger RL, et al. Definition and classification of cancer cachexia: an international consensus. Lancet Oncol 2011;12:489–95.
- [7] White JV, Guenter P, Jensen G, Malone A, Schofield M, Academy Malnutrition Work Group; A.S.P.E.N. Malnutrition Task Force; A.S.P.E.N. Board of Directors. Consensus statement: academy of nutrition and Dietetics and American society for parenteral and enteral nutrition: characteristics recommended for the identification and documentation of adult malnutrition (undernutrition). J Parenter Enter Nutr 2012;36:275–83.
- [8] Cederholm T, Bosaeus I, Barazzoni R, Bauer J, Van Gossum A, Klek S, et al. Diagnostic criteria for malnutrition — an ESPEN consensus statement. Clin Nutr 2015;34:335–40.
- [9] Jensen GL. Global leadership conversation: addressing malnutrition. J Parenter Enter Nutr 2016;40:455–7.
- [10] Cederholm T, Jensen GL. To create a consensus on malnutrition diagnostic criteria: a report from the Global Leadership Initiative on Malnutrition (GLIM) meeting at the ESPEN Congress 2016. Clin Nutr 2017;36:7–10.
- [11] Jensen GL, Cederholm T. Global leadership initiative on malnutrition: progress report from ASPEN clinical nutrition week 2017. J Parenter Enter Nutr 2017. https://doi.org/10.1177/0148607117707761.
- [12] Kondrup J, Allison SP, Elia M, Vellas B, Plauth M, Educational and Clinical Practice Committee, European Society of Parenteral and Enteral Nutrition (ESPEN). ESPEN guidelines for nutrition screening 2002. Clin Nutr 2003;22: 415–21
- [13] Skipper A, Ferguson M, Thompson K, Castellanos VH, Porcari J. Nutrition screening tools: an analysis of the evidence. J Parenter Enter Nutr 2012;36(3):
- [14] van Bokhorst-de van der Schueren MA, Guaitoli PR, Jansma EP, de Vet HC. Nutrition screening tools: does one size fit all? A systematic review of screening tools for the hospital setting. Clin Nutr 2014;33:39–58.

- [15] Cruz-Jentoft AJ, Baeyens JP, Bauer JM, Boirie Y, Cederholm T, Landi F, et al., European Working Group on Sarcopenia in Older People. Sarcopenia: European consensus on definition and diagnosis: report of the European working group on sarcopenia in older people. Age Ageing 2010;39:412–23.
- [16] Studenski SA, Peters KW, Alley DE, Cawthon PM, McLean RR, Harris TB, et al. The FNIH sarcopenia project: rationale, study description, conference recommendations, and final estimates. J Gerontol A Biol Sci Med Sci 2014;69: 547–58.
- [17] Chen LK, Lee WJ, Peng LN, Liu LK, Arai H, Akishita M, Asian working group for sarcopenia. Recent advances in sarcopenia research in Asia: updat from the Asian working group for sarcopenia. JAMDA 2016;17:767.e1–7.
 [18] Delmonico MJ, Harris TB, Visser M, Park SW, Conroy MB, Velasquez-Mieyer P,
- [18] Delmonico MJ, Harris TB, Visser M, Park SW, Conroy MB, Velasquez-Mieyer P, et al. Longitudinal study of muscle strength, quality, and adipose tissue infiltration. Am J Clin Nutr 2009;90:1579–85.
- [19] Jensen GL, Hsiao PY, Wheeler D. Adult nutrition assessment tutorial. J Parenter Enter Nutr 2012:36:267—74.
- [20] Fried LP, Tangen CM, Walston J, Newman AB, Hirsch C, Gottdiener J, et al., Cardiovascular Health Study Collaborative Research Group. Frailty in older adults: evidence for a phenotype. J Gerontol A Biol Sci Med Sci 2001;56: M146–56.
- [21] Rubenstein LZ, Harker JO, Salvà A, Guigoz Y, Vellas B. Screening for undernutrition in geriatric practice: developing the short-form mini-nutritional assessment (MNA-SF). J Gerontol A Biol Sci Med Sci 2001;56:M366-72.
- [22] Stratton RJ, Hackston A, Longmore D, Dixon R, Price S, Stroud M, et al. Malnutrition in hospital outpatients and inpatients: prevalence, concurrent validity and ease of use of the 'malnutrition universal screening tool' ('MUST') for adults. Br J Nutr 2004;92:799–808.
- [23] Fouque D, Kalantar-Zadeh K, Kopple J, Cano N, Chauveau P, Cuppari L, et al. A proposed nomenclature and diagnostic criteria for protein—energy wasting in acute and chronic kidney disease. Kidney Int 2008;73:391—8.
- [24] Baumgartner RN, Koehler KM, Gallagher D, Romero L, Heymsfield SB, Ross RR, et al. Epidemiology of sarcopenia among the elderly in New Mexico. Am J Epidemiol 1998;147:755–63.
- [25] Chiles Shaffer N, Ferrucci L, Shardell M, Simonsick EM, Studenski S. Agreement and predictive validity using less-conservative foundation for the National Institutes of health sarcopenia project weakness cut points. J Am Geriatr Soc 2017;65(3):574–9.