

Sarcopenia: qué es y dónde estamos

Alfonso J. Cruz-Jentoft

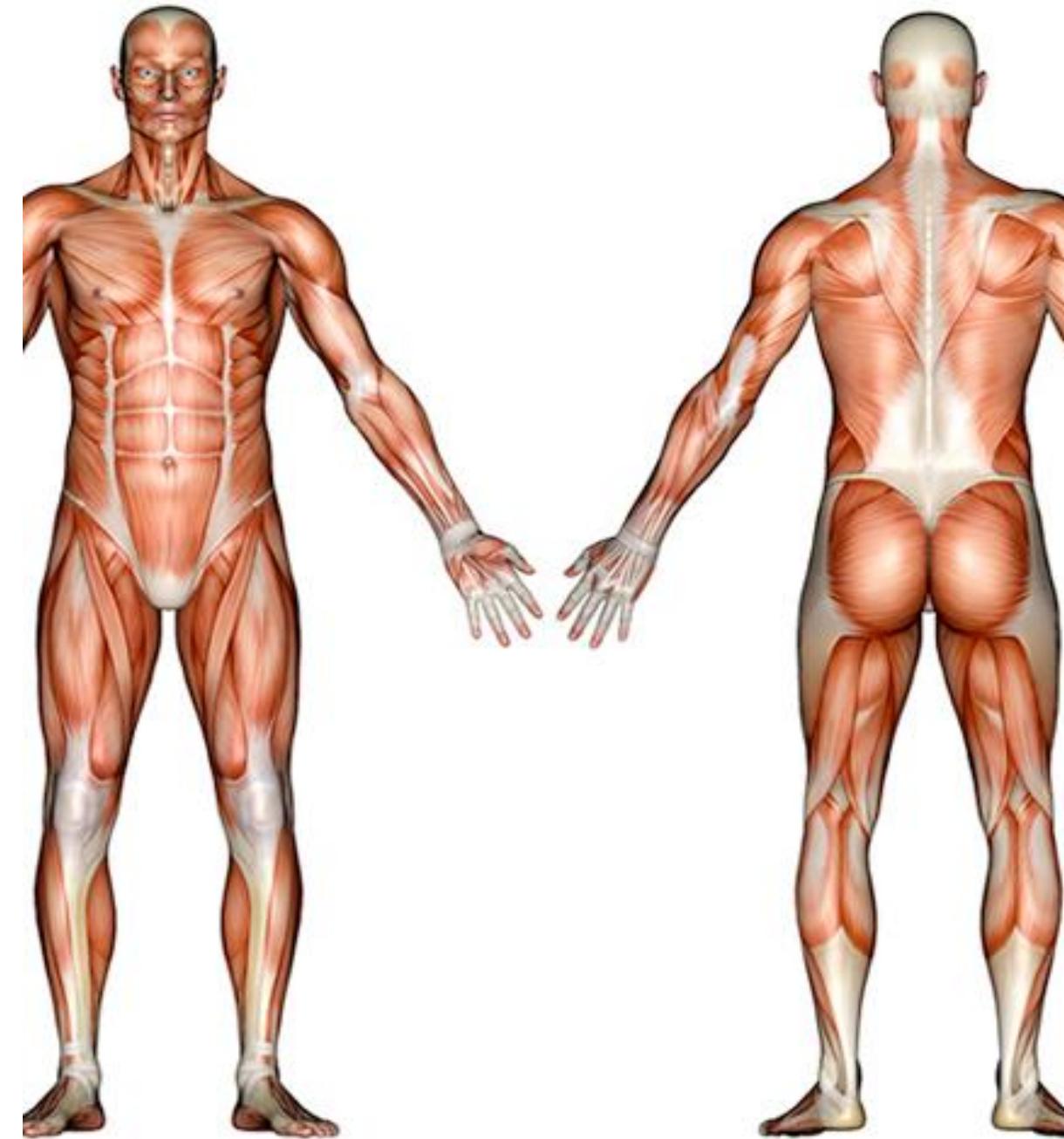
Servicio de Geriatría

Hospital Universitario Ramón y Cajal (IRYCIS),

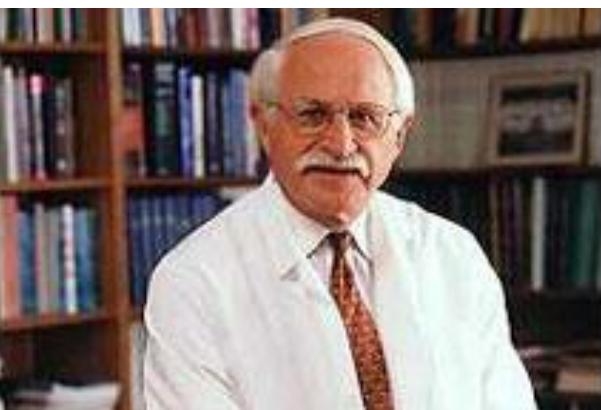
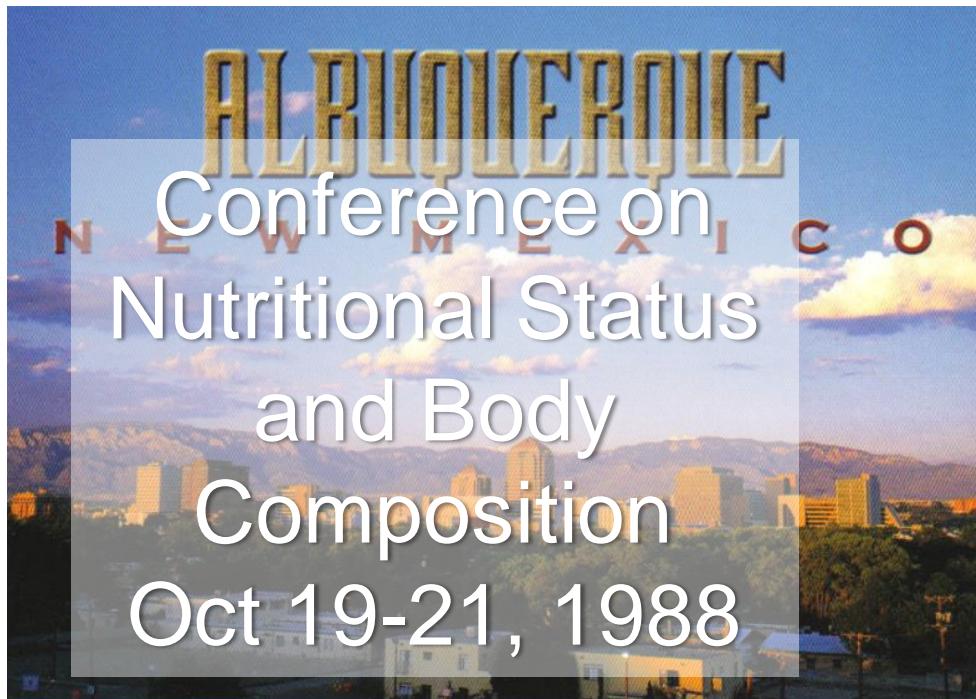
Madrid

El músculo humano: el órgano olvidado

- 600 músculos en el cuerpo humano
- Músculo esquelético: 40-45% de la masa corporal total
- 55% de este músculo está en las piernas
- 50% de las proteínas del cuerpo están en los músculos
 - movilidad
 - regulación de las proteínas
 - regulación endocrina



Sarcopenia: primeros pasos



- “No decline with age is as dramatic or potentially more significant than the decline in lean body mass. In fact, there may be no single feature of age-related decline more striking than the decline in lean body mass in affecting ambulation, mobility, energy intake, overall nutrient intake and status, independence and breathing.”
- “I suggested that if this phenomenon were to be taken seriously, we had to give it a name. I proposed that the name for this phenomenon should be derived from the Greek.”

Sarcopenia: primer acto científico

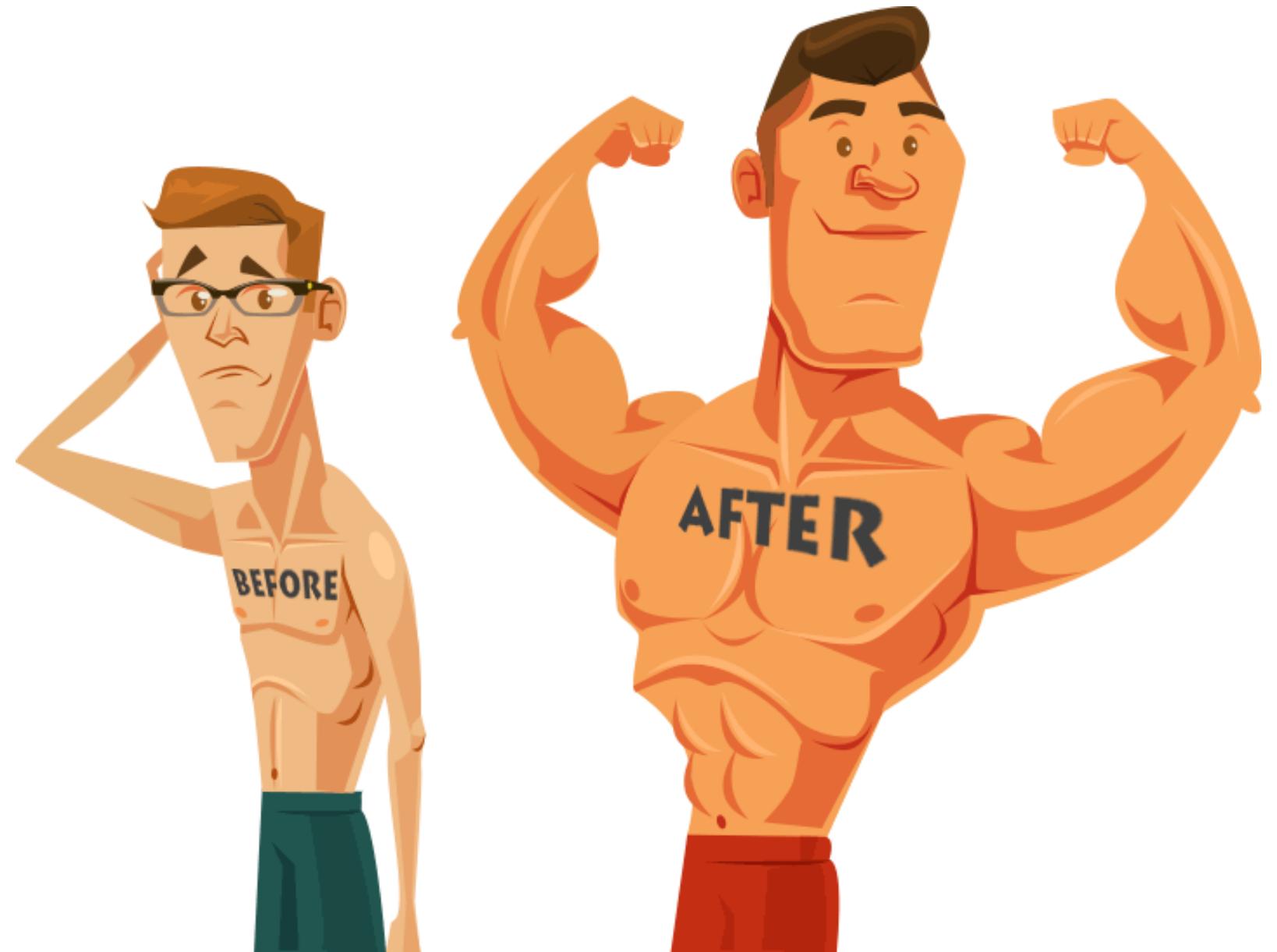
1st Workshop on Sarcopenia - NIA
September 19-21, 1994

"Sarcopenia" es un nombre genérico para la pérdida de masa, calidad y fuerza del músculo esquelético, que puede causar fragilidad en las personas mayores.

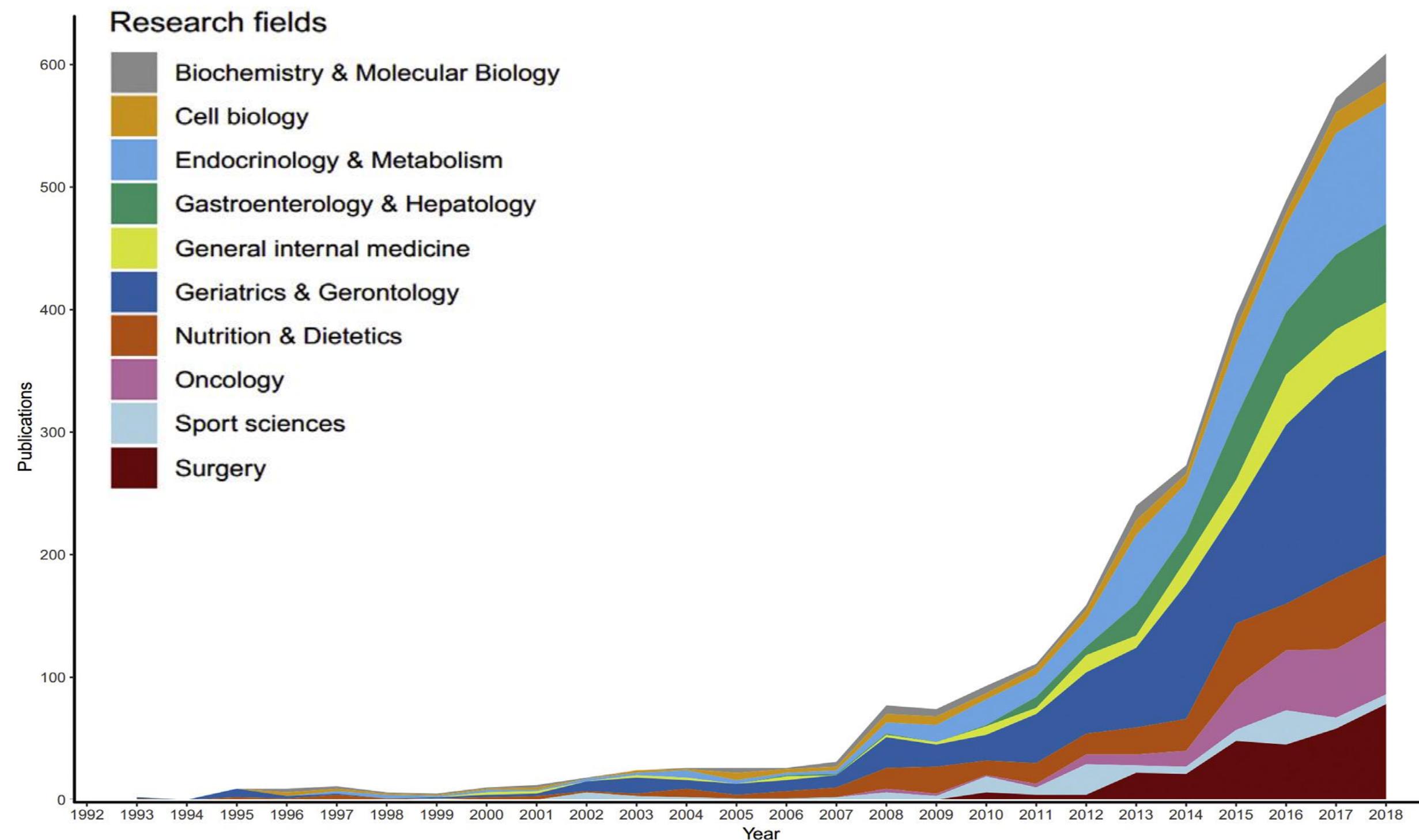
2010

Este año se da un
importante cambio en el
concepto de sarcopenia

Sarcopenia
≠
Baja masa
muscular

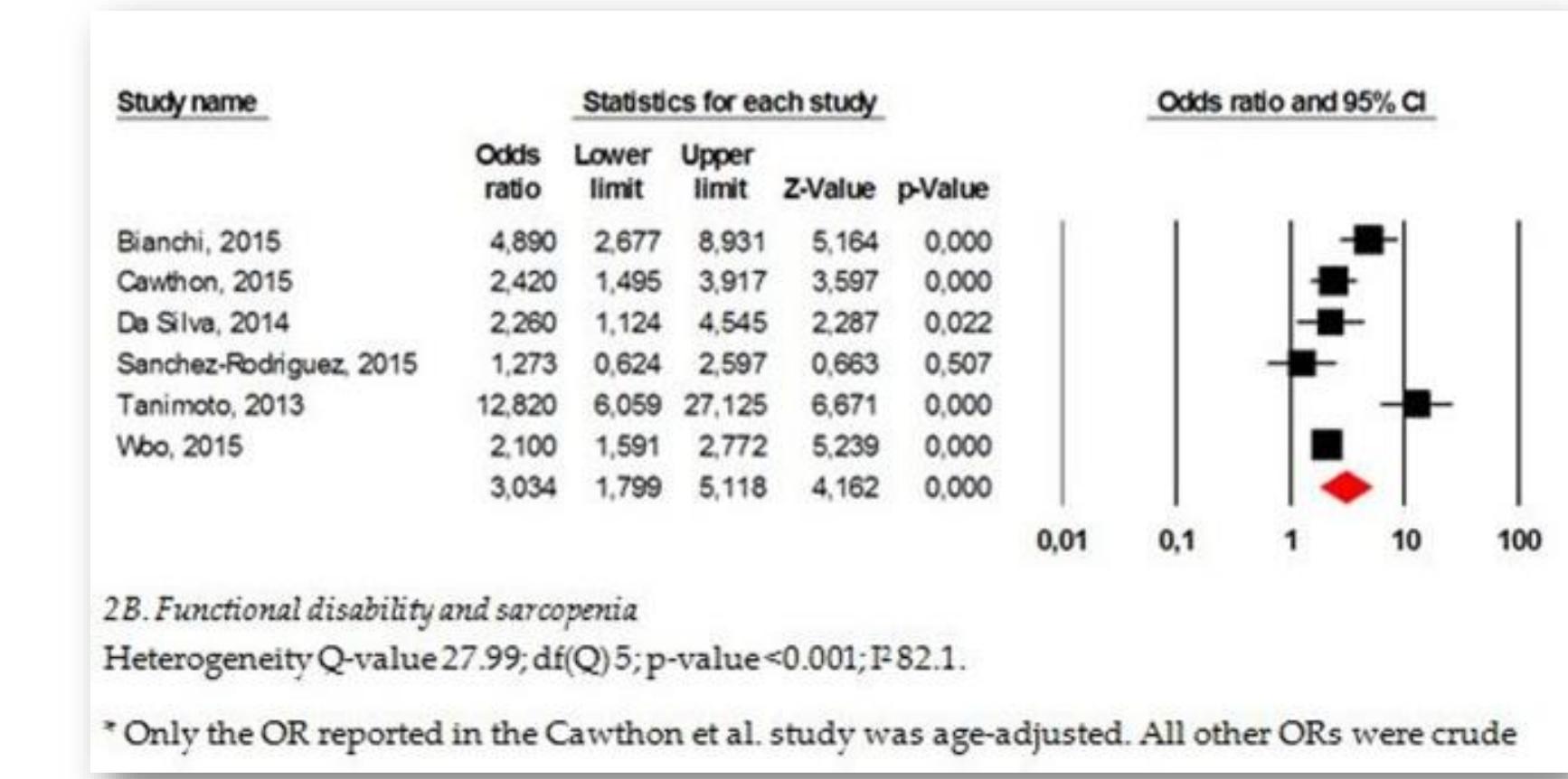
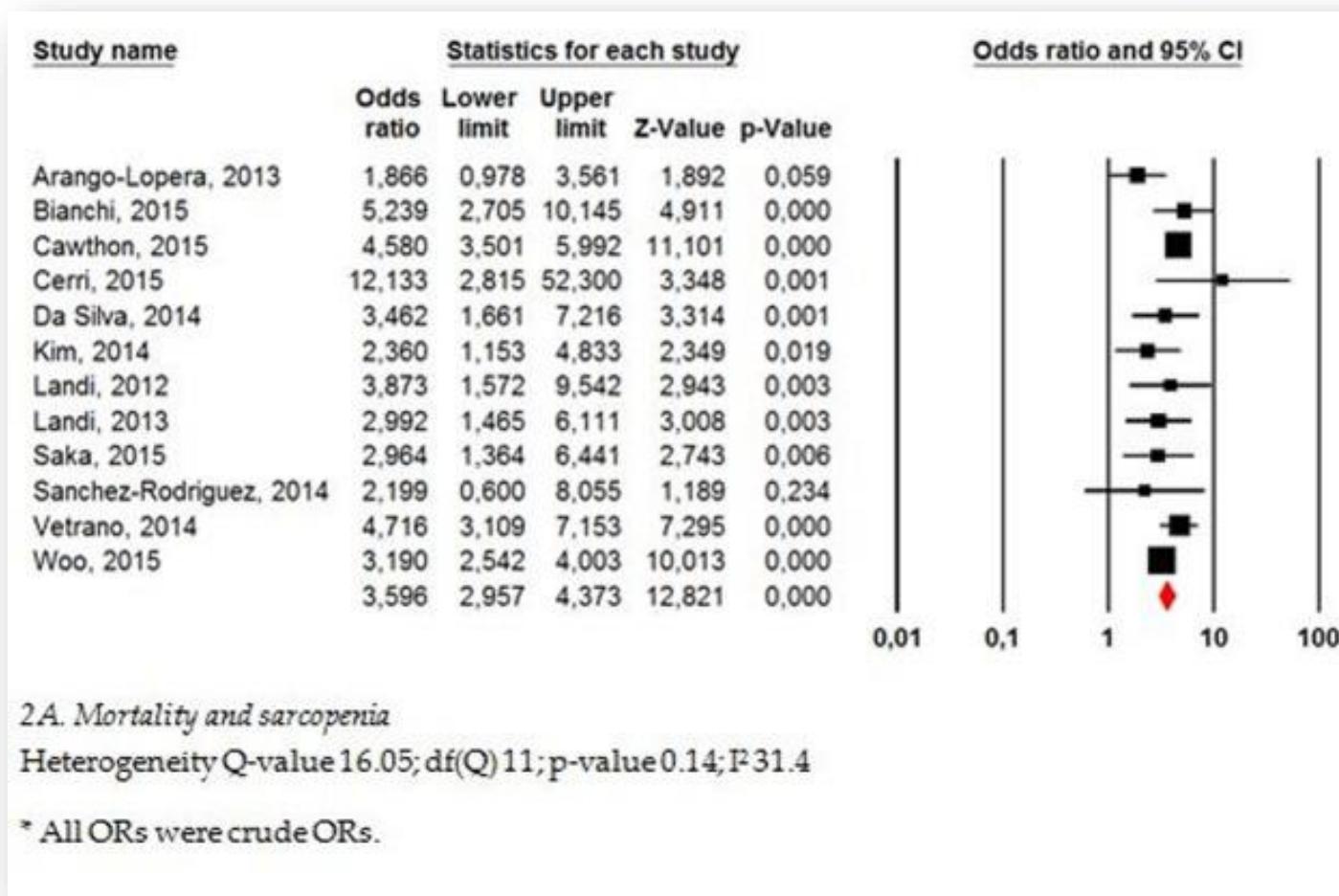


Crecimiento en la investigación



Yang M, Tan L, Li W. Landscape of Sarcopenia Research (1989–2018): A Bibliometric Analysis. *J Am Med Dir Assoc.* 2020 Mar;21(3):436-437.

Consecuencias de la sarcopenia



La sarcopenia sale del ámbito de la geriatría

THE LANCET

Seminar



Sarcopenia

Alfonso J Cruz-Jentoft, Avan A Sayer

Lancet 2019; 393: 2636-46

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Servicio de Geriatría, Hospital Universitario Ramón y Cajal (IRYCIS). Madrid, Spain

Sarcopenia is a progressive and generalised skeletal muscle disorder involving the accelerated loss of muscle mass and function that is associated with increased adverse outcomes including falls, functional decline, frailty, and mortality. It occurs commonly as an age-related process in older people, influenced not only by contemporaneous risk factors, but also by genetic and lifestyle factors operating across the life course. It can also occur in mid-life in association with a range of conditions. Sarcopenia has become the focus of intense research aiming to translate current knowledge about its pathophysiology into improved diagnosis and treatment, with particular interest in the development of biomarkers, nutritional interventions, and drugs to augment the beneficial effects of resistance exercise. Designing effective preventive strategies that people can apply during their lifetime is of primary concern. Diagnosis, treatment, and prevention of sarcopenia is likely to become part of routine clinical practice.

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GUIDELINES

Sarcopenia on definition

ALFONSO J. CRUZ-JE
TOMMY CEDERHOLM
STÉPHANE M. SCHNI
MARJOLEIN VISSER¹⁵,
SARCOPENIA IN OLD

ESTABLISHING IN AUSTRALIA AND

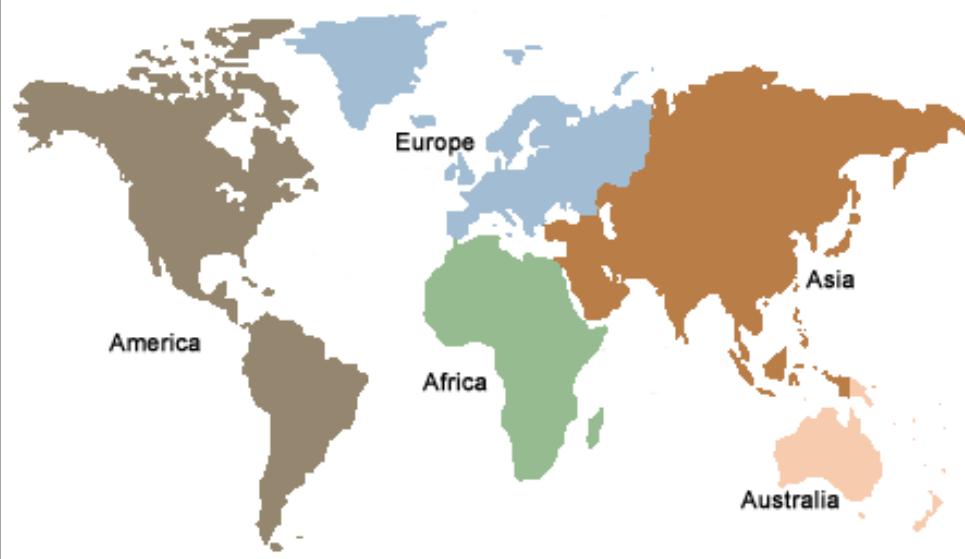
J. ZANKER^{1,2}, D. SCOTT^{1,2},
M. GROSSMANN⁷, A. HA
J.W.L. KEOUGH¹⁵⁻¹⁷, J.R
M. SIM¹⁸, R. VISVANATI
AUSTRALIAN AND NE
TASK F



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Liang-Kung Chen MD,
Tung-Wai Auyeung MD,
Hak Chul Jang MD,
Taro Kojima MD, PI
Sang Yoon Lee MD,
Chih-Kuang Liang MD,
Li-Ning Peng MD, N
Chang Won Won MD,
Masahiro Akishita I

SPECIAL ARTICLE

Sarcopenia Definition: The Position Statements of the Sarcopenia Definition and Outcomes Consortium

Shalender Bhagat, MB, BS, * Thomas G. Travison, PhD, † Todd M. Manini, PhD, ‡
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Greg Gorsicki, PhD, §§§§§ Rosaly Correa-De-Araujo, MD, PhD, §§§§§§ and Peggy M. Cawthon, PhD §

GUIDELINES

Sarcopenia: revised European consensus on definition and diagnosis

ALFONSO J. CRUZ-JENTOFF¹, GÜLISTAN BAHAT², JÜRGEN BAUER³, YVES BOIRIE⁴, OLIVIER BRUYÈRE⁵, TOMMY CEDERHOLM⁶, CYRUS COOPER⁷, FRANCESCO LANDI⁸, YVES ROLLAND⁹, AVAN AIHIE SAYER¹⁰, STÉPHANE M. SCHNEIDER¹¹, CORNEL C. SIEBER¹², EVA TOPINKOVA¹³, MAURITS VANDEWOODE¹⁴, MARJOLEIN VISSER¹⁵, MAURO ZAMBONI¹⁶, WRITING GROUP FOR THE EUROPEAN WORKING GROUP ON SARCOPENIA IN OLDER PEOPLE 2 (EWGSOP2), AND THE EXTENDED GROUP FOR EWGSOP2

EWGSOP2

Definición de sarcopenia

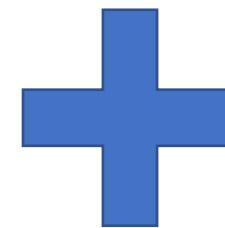
- La sarcopenia es una enfermedad progresiva y generalizada del músculo esquelético que se asocia con un aumento del riesgo de sufrir consecuencias adversas como fracturas, incapacidad física y mortalidad.

Insuficiencia muscular

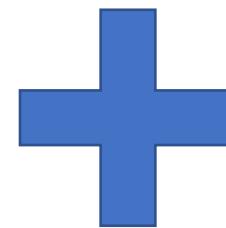
EWGSOP²

Definición operativa de sarcopenia

BAJA FUERZA
MUSCULAR



BAJA MASA/
CALIDAD
MUSCULAR



BAJO RENDIMIENTO
FÍSICO

**SARCOPENIA
PROBABLE**

**SARCOPENIA
CIERTA**

**SARCOPENIA
GRAVE**

EWGSOP²

Detección de casos de sarcopenia

- En la tercera edad más tarde la sarcopenia
- El EGSOP2 detecta los casos

Preguntas		Puntaje
Strength (Fuerza)	¿Qué tanta dificultad tiene para llevar o cargar 4.5 kg?	Ninguna = 0 Alguna = 1 Mucho o incapaz = 2
Assistance in walking (Asistencia para caminar)	¿Qué tanta dificultad tiene para cruzar caminando por un cuarto?	Ninguna = 0 Alguna = 1 Mucho, usando auxiliares o incapaz = 2
Rise from chair (Levantarse de una silla)	¿Qué tanta dificultad tiene para levantarse de una silla o cama?	Ninguna = 0 Alguna = 1 Mucho o incapaz sin ayuda = 2
Climb stairs (Subir escaleras)	¿Qué tanta dificultad tiene para subir 10 escalones?	Ninguna = 0 Alguna = 1 Mucho o incapaz = 2
Falls (Caídas)	¿Cuántas veces se ha caído en el último año?	Ninguna = 0 1-3 caídas = 1 4 o más caídas = 2

Puntuación total: _____

Interpretación

Alta probabilidad de sarcopenia = 4 o más probabilidades.
1, 2 ó 3 puntos = Baja probabilidad de sarcopenia.

En la tercera edad más tarde la sarcopenia cuando el paciente siente sensación de debilidad, fatiga o pérdida de energía para hacer pruebas para

SARC-F para detectar



Puntos de corte

Test	Puntos de corte en hombres	Puntos de corte en mujeres	Referencias
Puntos de corte EWGSOP2 para definir la baja fuerza muscular			
Fuerza de prensión	<27 kg	<16 kg	Dodds, 2014[26]
Levantarse de la silla	>15 sec for 5 rises		Cesari, 2009[67]
Puntos de corte EWGSOP2 para definir la baja masa muscular			
ASM	< 20 kg	< 15 kg	Studenski, 2014[3]
ASM/height ²	< 7.0 kg/m ²	< 5.5 kg/m ²	Gould, 2014[125]

Puntos de corte

Test	Puntos de corte en hombres	Puntos de corte en mujeres	Referencias
Puntos de corte EWGSOP2 para definir el bajo rendimiento físico			
Velocidad de marcha		$\leq 0.8 \text{ m/s}$	Cruz-Jentoft, 2010[1] Studenski, 2011[84]
SPPB		$\leq 8 \text{ puntos}$	Pavasini, 2016[90] Guralnik, 1995[126]
TUG		$\geq 20 \text{ s}$	Bischoff, 2003[127]
400m walk test		No completarlo o tardar $\geq 6 \text{ min}$ en hacerlo	Newman, 2006[128]

P2

Sarcopenia aguda y crónica

- Sarcopenia aguda: menos de 6 meses
 - Normalmente asociada a enfermedad o lesión aguda
- Sarcopenia crónica: dura \geq 6 meses.
 - Asociada a enfermedades crónicas o progresivas
- Importancia de medir la sarcopenia a lo largo del tiempo.
- Puede facilitar la intervención precoz.

EWGSOP2

Sarcopenia aguda

Table 1. Factors and diagnostic tools concerning the diagnosis of acute sarcopenia

General aspects		
<i>Patient factors</i> Health conditions (pain, dyspnoea, delirium/dementia, reduced mobility/bed bound), intravenous lines, catheters, and patient refusal		
<i>Staff/hospital factors</i> Lack of time and staff, lack of knowledge, and no availability of equipment		
Measurements		
Muscle mass	DXA CT MRI BIA Ultrasound	DXA, CT, MRI: – Expensive, exposure to radiation, subject to availability, and patient's clinical status to be transferred to the facility DXA and BIA: – Influenced by hydration situation Ultrasound: – Nonuniversal cutoff points
Muscle strength	Hand dynamometer	Influenced by mental situation and uncontrolled pain
Physical performance	SPPB TUG Gait speed	Influenced by acute illness, comorbidities, and uncontrolled pain

Categorías de la sarcopenia



Nuevo algoritmo

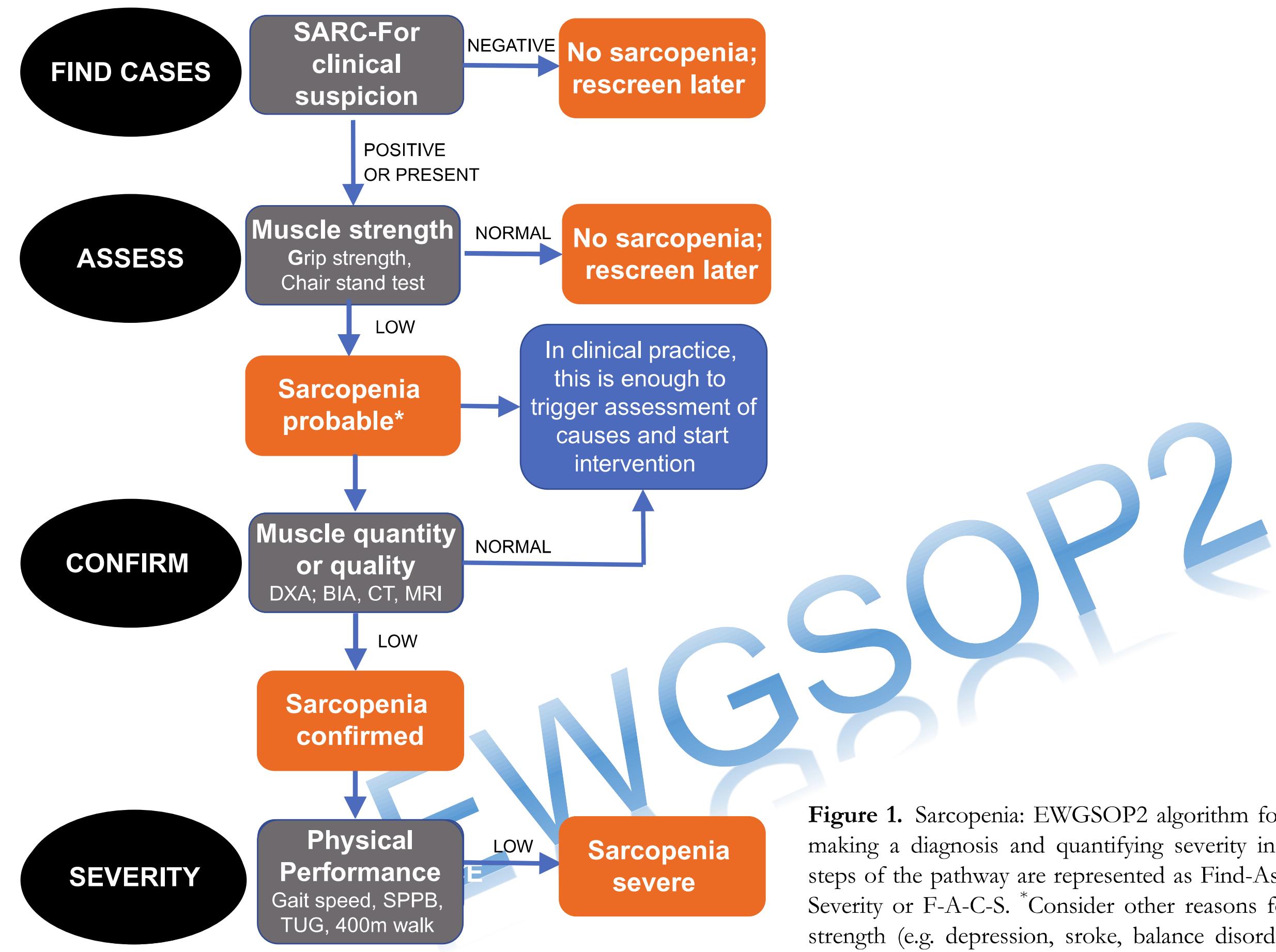


Figure 1. Sarcopenia: EWGSOP2 algorithm for case-finding, making a diagnosis and quantifying severity in practice. The steps of the pathway are represented as Find-Assess-Confirm-Severity or F-A-C-S. *Consider other reasons for low muscle strength (e.g. depression, stroke, balance disorders, peripheral vascular disorders).

ESPEN / EASO definition of sarcopenic obesity

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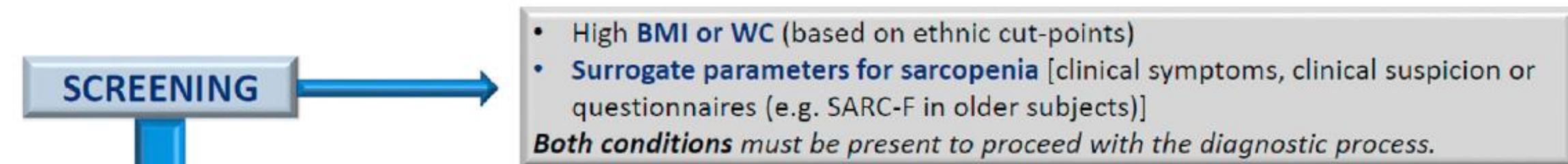
ESPEN Guideline

Definition and diagnostic criteria for sarcopenic obesity: ESPEN and EASO consensus statement[☆]

Lorenzo M. Donini ^{a,*}, Luca Busetto ^b, Stephan C. Bischoff ^c, Tommy Cederholm ^d, Maria D. Ballesteros-Pomar ^e, John A. Batsis ^f, Juergen M. Bauer ^g, Yves Boirie ^h, Alfonso J. Cruz-Jentoft ⁱ, Dror Dicker ^j, Stefano Frara ^k, Gema Frühbeck ^l, Laurence Genton ^m, Yftach Gepner ⁿ, Andrea Giustina ^k, Maria Cristina Gonzalez ^o, Ho-Seong Han ^p, Steven B. Heymsfield ^q, Takashi Higashiguchi ^r, Alessandro Laviano ^a, Andrea Lenzi ^a, Ibolya Nyulasi ^s, Edda Parrinello ^a, Eleonora Poggiogalle ^a, Carla M. Prado ^t, Javier Salvador ^u, Yves Rolland ^v, Ferruccio Santini ^w, Mireille J. Serlie ^x, Hanping Shi ^y, Cornel C. Sieber ^z, Mario Siervo ^{aa}, Roberto Vettor ^b, Dennis T. Villareal ^{ab}, Dorothee Volkert ^z, Jianchun Yu ^{ac}, Mauro Zamboni ^{ad}, Rocco Barazzoni ^{ae,**}



ESPEN / EASO definition of sarcopenic obesity



¿Qué sucede en la práctica clínica real?

177 organizaciones
en Reino Unido

104 (59%) responden
sobre fragilidad

49 (28%) responden
sobre sarcopenia

26 (53%) afirman
que identifican la
sarcopenia

19 (39%) usan alguna
herramienta
diagnóstica

3 (6%) usan criterios
diagnósticos

Muscle mass (%)	
Bioimpedance assessment	2 (11)
Dual X-ray absorptiometry	3 (16)
Computed tomography	2 (11)
Magnetic resonance imaging	1 (5)
Observation or anthropometry (%)	7 (37)
Muscle function (%)	
Walk speed	16 (84)
Grip strength	10 (53)
Questionnaire / history (%)	
Number measuring muscle mass (%)	4 (21)
Number measuring muscle function (%)	18 (95)
Number measuring muscle mass AND muscle function (%)	4 (21)

Table 1. Tools used to identify sarcopenia (n=19 organisations).

¿Qué sucede en la práctica clínica real?

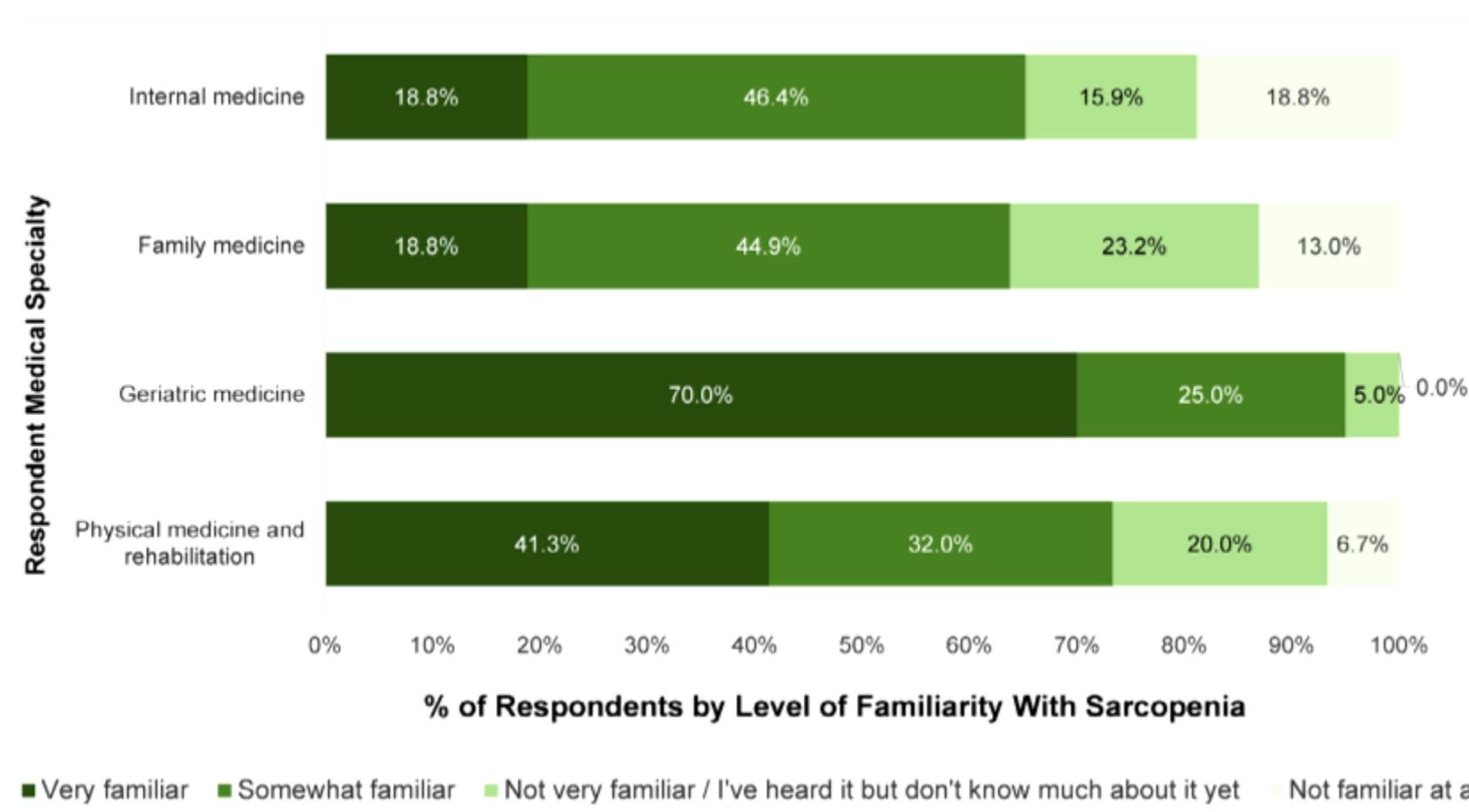


TABLE 1 Physicians' diagnostic techniques and diagnostic criteria for sarcopenia

Measurements used in diagnosis (select all that apply; n = 248)	Percent
Manual muscle testing strength scale	40%
Gait speed test results	39%
Grip strength test results (e.g., from a dynamometer)	35%
None-I do not use specific measurements to confirm a diagnosis.	21%
DEXA scan results	19%
Electrodiagnostic results	14%
CT/MRI results	10%
Ultrasound results	5%
Other	7%
N/A-I always defer to a specialist for diagnosis	1%
Diagnostic criteria (select one; n = 246)	
N/A-I do not typically use specific diagnostic criteria.	50%
Unsure which one applies to my criteria	27%
FNIH (Foundation for the National Institutes of Health)	14%
SSCWD (Society of Sarcopenia, Cachexia, and Wasting Disorders)	6%
EWGSOP (European Working Group on Sarcopenia in Older People)	4%
Other	0%

GUIDELINES

Sarcopenia on definition

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MARJOLEIN VISSER¹⁵,
SARCOPENIA IN OLD

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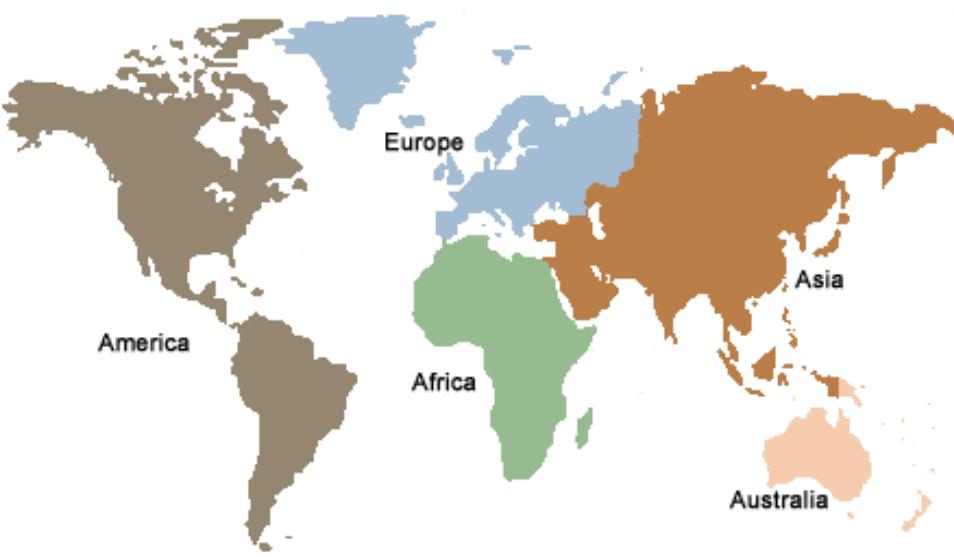
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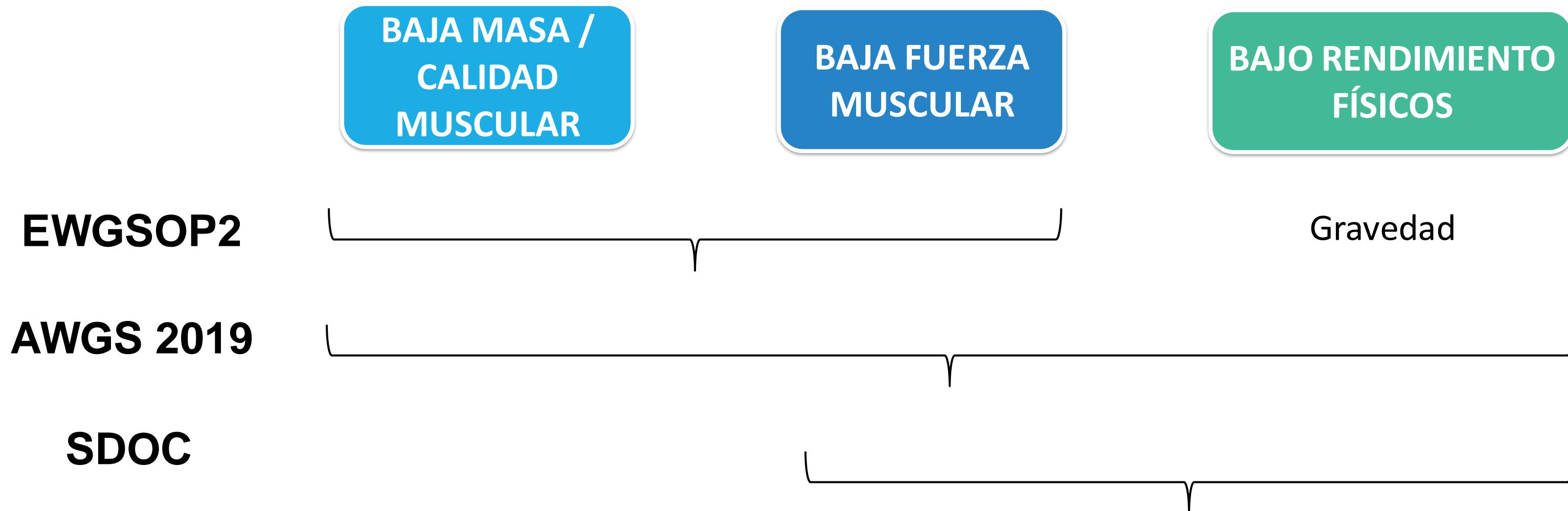
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SPECIAL ARTICLE

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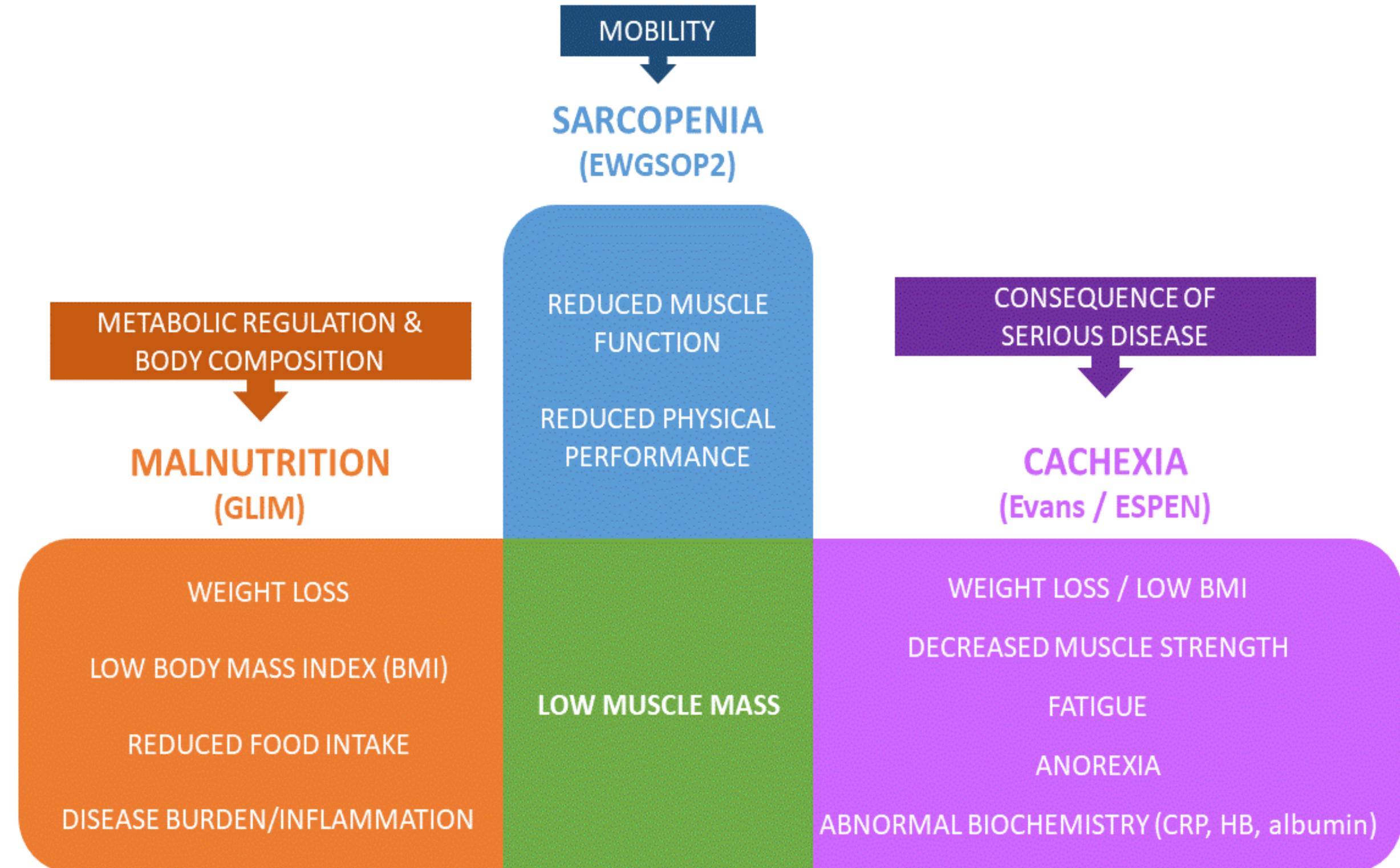
¿Qué elementos tienen la definiciones modernas de sarcopenia?



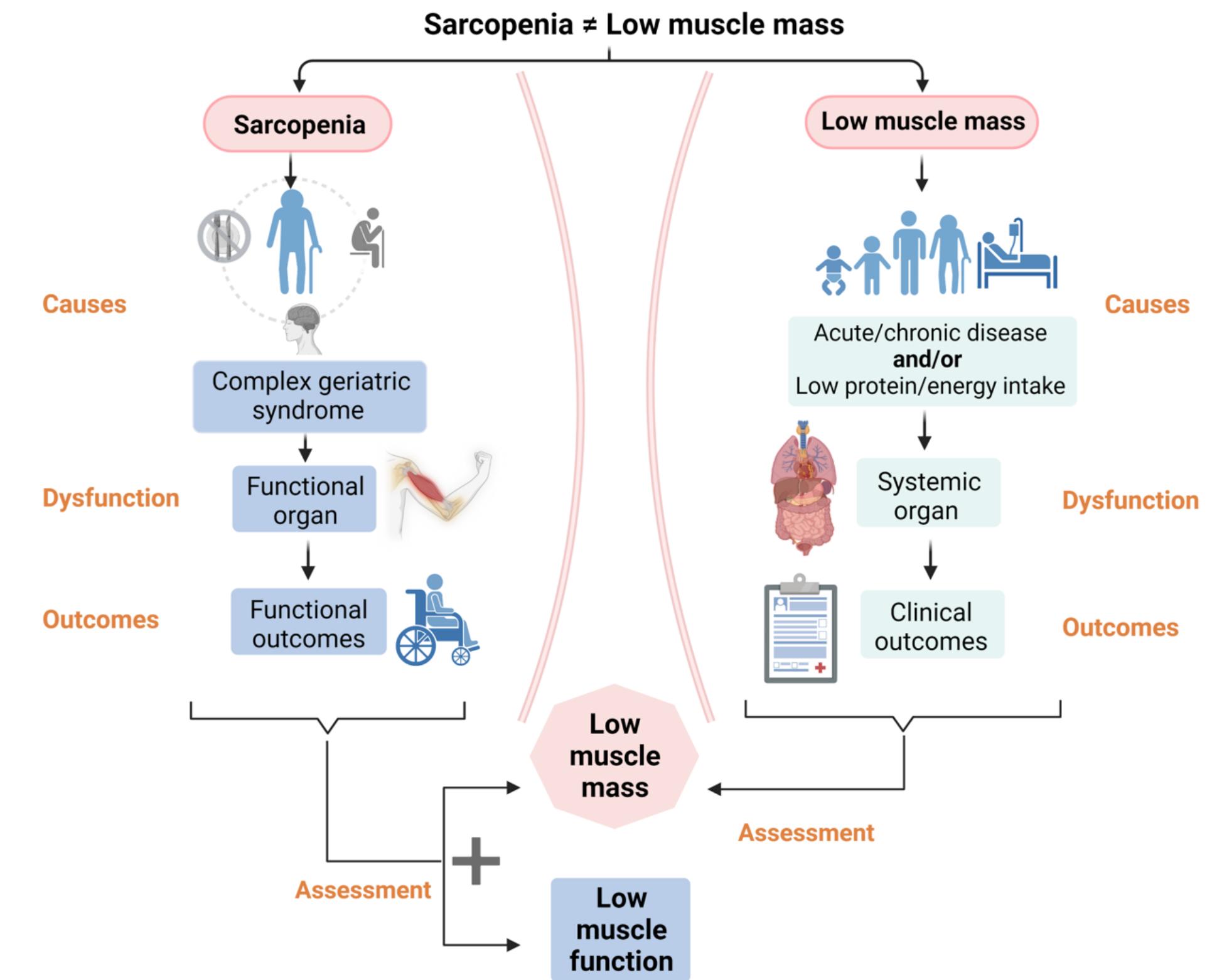
Definition	Muscle mass	Muscle strength	Physical performance	Cut-offs defined	Comment
EWGSOP (2010)	●	●	●		Insufficient evidence to propose cut-offs
IWGS (2011)	●		●	●	
SSCWD (2011)	●		●	●	Named “sarcopenia with limited mobility”
AWGS (2014)	●	●	●	●	Same as EWGSOP but adds cut-offs for Asia
FNIH (2014)	●	●		●	Uses physical performance as outcome
EWGSOP2 (2019)	●	●	●	●	Physical performance used for severity
AWGS (2019)	●	●	●	●	
SDOC (2019)		●	●	●	

Adapted from Cruz-Jentoft AJ, Sayer AA. Sarcopenia. Lancet. 2019;393(10191):2636-2646.

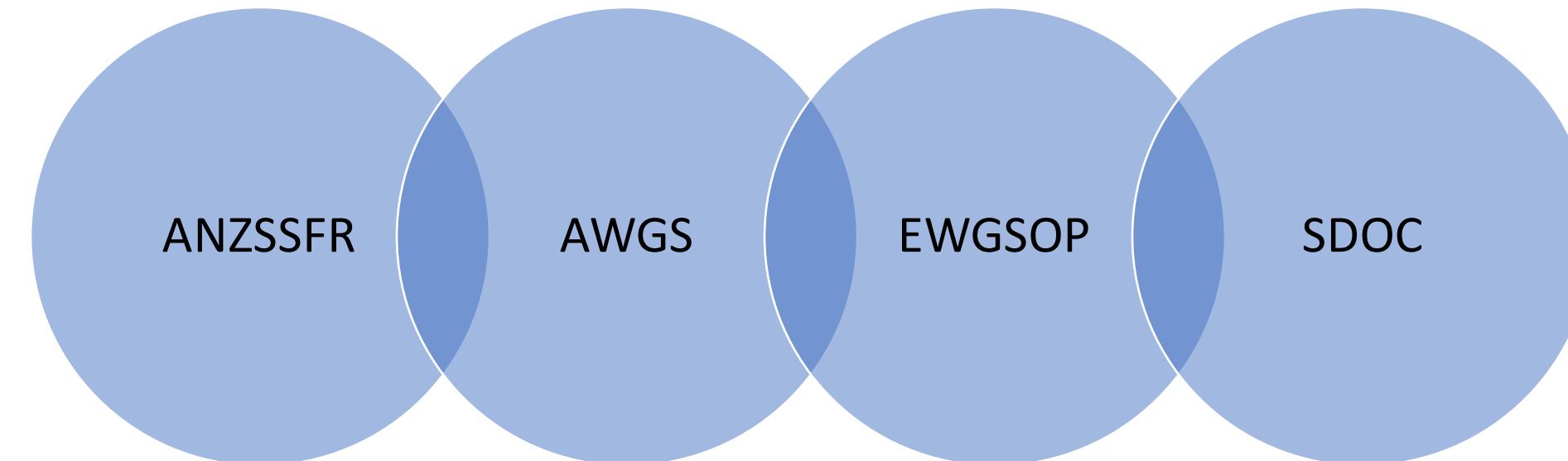
La masa muscular en la encrucijada



Sarcopenia ≠ Low muscle mass



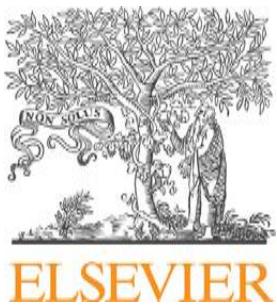
The GLIS initiative



Australian and New Zealand Society
for Sarcopenia and Frailty Research



¿Por qué GLIS?



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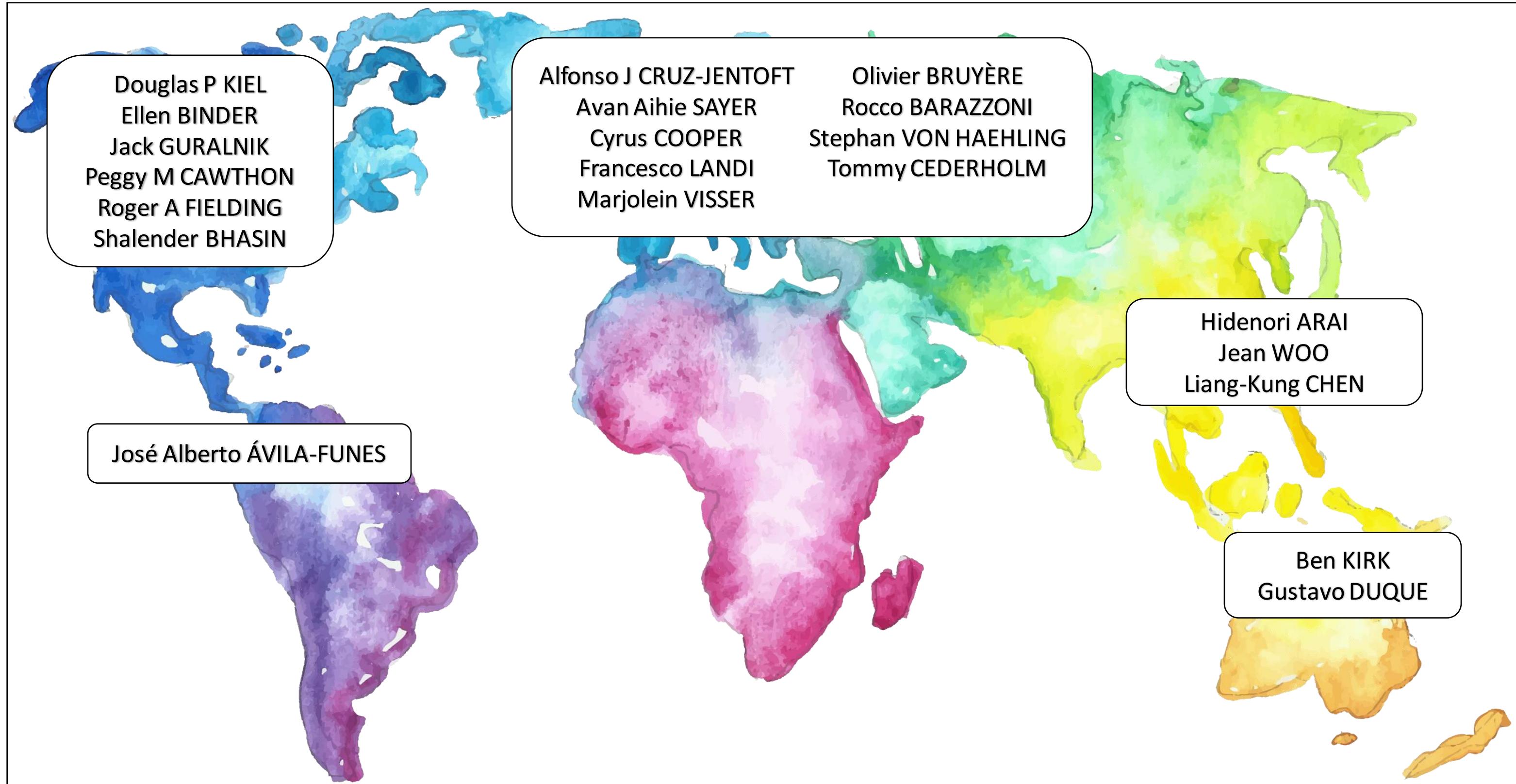


ESPEN Endorsed Recommendation

GLIM criteria for the diagnosis of malnutrition – A consensus report from the global clinical nutrition community[☆]

T. Cederholm ^{a, b, *, 1}, G.L. Jensen ^{c, 1}, M.I.T.D. Correia ^d, M.C. Gonzalez ^e, R. Fukushima ^f,
T. Higashiguchi ^g, G. Baptista ^h, R. Barazzoni ⁱ, R. Blaauw ^j, A. Coats ^{k, l}, A. Crivelli ^m,
D.C. Evans ⁿ, L. Gramlich ^o, V. Fuchs-Tarlovsky ^p, H. Keller ^q, L. Llido ^r, A. Malone ^{s, t},
K.M. Mogensen ^u, J.E. Morley ^v, M. Muscaritoli ^w, I. Nyulasi ^x, M. Pirlich ^y, V. Pisprasert ^z,
M.A.E. de van der Schueren ^{aa, ab}, S. Siltharm ^{ac}, P. Singer ^{ad, ae}, K. Tappenden ^{af},
N. Velasco ^{ag}, D. Waitzberg ^{ah}, P. Yamwong ^{ai}, J. Yu ^{aj}, A. Van Gossum ^{ak, 2}, C. Compher ^{al, 2},
GLIM Core Leadership Committee, GLIM Working Group³

GLIS steering committee



SPECIAL ARTICLE



Defining terms commonly used in sarcopenia research: a glossary proposed by the Global Leadership in Sarcopenia (GLIS) Steering Committee

Peggy M. Cawthon^{1,2} · Marjolein Visser^{3,4} · Hidenori Arai⁵ · José A. Ávila-Funes⁶ · Rocco Barazzoni⁷ · Shalender Bhasin⁸ · Ellen Binder^{9,10} · Olivier Bruyère¹¹ · Tommy Cederholm^{12,13} · Liang-Kung Chen^{14,15,16} · Cyrus Cooper^{17,18} · Gustavo Duque^{19,20} · Roger A. Fielding²¹ · Jack Guralnik²² · Douglas P. Kiel²³ · Ben Kirk^{24,25} · Francesco Landi²⁶ · Avan A. Sayer²⁷ · Stephan Von Haehling^{28,29} · Jean Woo^{30,31} · Alfonso J. Cruz-Jentoft³²

Glosario



Muscle density

A measure of the attenuation of X-rays through the muscle tissue expressed in Hounsfield units (HU).

Muscle texture

A measure of the spatial arrangement, contrast, and consistency of the greyscale pixels in the image.



Muscle specific force

Strength standardized to muscle size

GLIS: metodología

Método **Delphi** con escala Likert de 11 puntos, desde 0 (totalmente en desacuerdo) a 10 (totalmente de acuerdo)

107 participantes (edad media 54 años , 64% hombres) de **29 países** y 7 continentes/regiones.

GLIS: definición conceptual de sarcopenia

La sarcopenia es una enfermedad generalizada del **músculo esquelético**.

Su prevalencia aumenta con la **edad**.

La definición conceptual **no cambia** con el lugar de atención (p.ej. hospitalizados vs. pacientes externos), edad o comorbilidad (p. ej. ICC, ERC, cancer) y es la misma en clínica que en investigación.

La sarcopenia is potencialmente **reversible**.

Componentes de la sarcopenia:

Masa muscular

Fuerza muscular

Fuerza muscular específica (fuerza/tamaño del músculo)

No hubo acuerdo en definir niveles de gravedad de la sarcopenia.

GLIS: definición conceptual de sarcopenia

Consecuencias de la sarcopenia:

- Empeoramiento del **rendimiento físico**
- Limitaciones de **movilidad** (caminar, transferencias)
- **Caídas** y fracturas
- Incapacidad para hacer **AVD básicas e instrumentales**
- **Hospitalizaciones**, ingresos en residencias
- Mala **calidad de vida**
- Mayor **mortalidad**

Sarcopenia: guía clínica

INTERNATIONAL CLINICAL PRACTICE GUIDELINES FOR SARCOPENIA (ICFSR): SCREENING, DIAGNOSIS AND MANAGEMENT

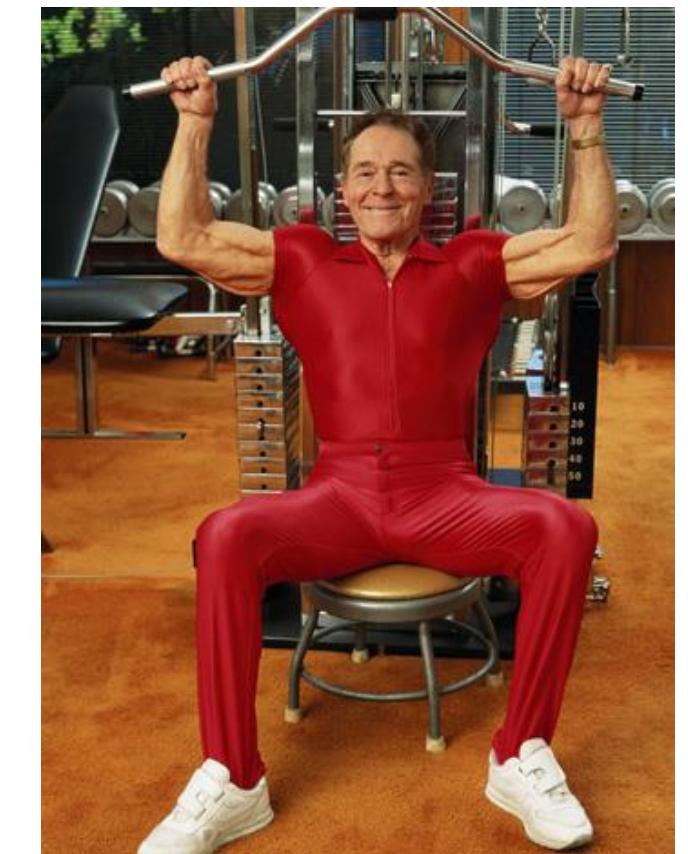
E. DENT^{1,2}, J.E. MORLEY³, A.J. CRUZ-JENTOFT⁴, H. ARAI⁵, S.B. KRITCHEVSKY⁶, J. GURALNIK⁷,
J.M. BAUER⁸, M. PAHOR⁹, B.C. CLARK¹⁰, M. CESARI^{11,12}, J. RUIZ¹³, C.C. SIEBER¹⁴, M. AUBERTIN-
LEHEUDRE¹⁵, D.L. WATERS¹⁶, R. VISVANATHAN¹⁷, F. LANDI¹⁸, D.T. VILLAREAL¹⁹, R. FIELDING²⁰,
C.W. WON²¹, O. THEOU^{17,22}, F.C. MARTIN²³, B. DONG²⁴, J. WOO²⁵, L. FLICKER²⁶, L. FERRUCCI²⁷,
R.A. MERCHANT²⁸, L. CAO²⁹, T. CEDERHOLM³⁰, S.M.L. RIBEIRO³¹, L. RODRÍGUEZ-MAÑAS³²,
S.D. ANKER^{33,34}, J. LUNDY³⁵, L.M. GUTIÉRREZ ROBLEDO³⁶, I. BAUTMANS^{37,38,39}, I. APRAHAMIAN⁴⁰,
J.M.G.A. SCHOLS⁴¹, M. IZQUIERDO⁴², B. VELLAS⁴³

Prevención y tratamiento de la sarcopenia

3. Physical Activity	3A. In patients with sarcopenia, prescription of resistance-based training may be effective to improve lean mass, strength and physical function	Strong	+++
4. Protein	4A. We recommend clinicians consider protein supplementation/a protein-rich diet for older adults with sarcopenia	Conditional	++
	4B. Clinicians may also consider discussing with patients the importance of adequate calorie and protein intake	Conditional	+
	4C. Nutritional (protein) intervention should be combined with a physical activity intervention	Conditional	++
5. Vitamin D	5A. Insufficient evidence exists to determine whether a Vitamin D supplementation regime by itself is effective in older adults with sarcopenia	Insufficient evidence	+
6. Anabolic Hormones	6A. The current evidence is insufficient to recommend anabolic hormones for the management of sarcopenia	Insufficient evidence	+
7. Pharmacologic Interventions	7A. Pharmacological interventions are not recommended as first-line therapy for the management of sarcopenia	Insufficient evidence	+
8. Research	8A.. Future international collaboration and large-scale RCTs focusing specifically on older people with sarcopenia are recommended	n/a	n/a

Definición de actividad física y ejercicio

Actividad física: cualquier movimiento del cuerpo producido por músculos esqueléticos que consume energía (kcal). En la vida diaria puede distinguirse la ocupacional, deportiva, doméstica y otras.



Ejercicio: un tipo de actividad física planificado, estructurado y repetitivo que busca mejorar o mantener el estado de forma física.

Low physical activity
No exercise

Low physical activity
Exercise

High physical activity
No exercise

High physical activity
Exercise

Ejercicio de resistencia y parámetros musculares

Fuerza muscular (fuerza máxima): capacidad de un grupo muscular de ejercer una fuerza máxima contra resistencia en una única contracción (no dependiente del tiempo).

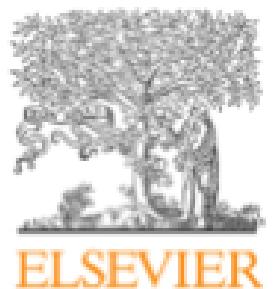
- Ej: levantar una vez un objeto pesado.
- Entrenamiento: movimientos lentos repetidos contra resistencia.

Potencia muscular (explosividad): gran producción de fuerza en un periodo de tiempo corto (dependiente del tiempo).

- Ej: salto vertical, patadas.
- Entrenamiento: repeticiones rápidas contra resistencia.

Durabilidad muscular (repeticiones): se mantiene una fuerza submáxima en un periodo de tiempo largo.

- Ej: saltos, bicicleta, natación.
- Entrenamiento: ejercicio aeróbico que estimula el consumo de energía.



JAMDA

journal homepage: www.jamda.com



Special Article

Evidence-based Recommendations for Optimal Dietary Protein Intake in Older People: A Position Paper From the PROT-AGE Study Group

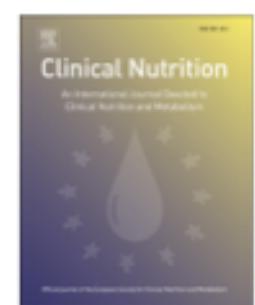
Jürgen Bauer MD^{a,*}, Gianni Biolo MD, PhD^b, Tommy Cederholm MD, PhD^c, Matteo Cesari MD, PhD^d, Alfonso J. Cruz-Jentoft MD^e, John E. Morley MB, BCh^f, Stuart Phillips PhD^g, Cornel Sieber MD, PhD^h, Peter Stehle MD, PhDⁱ, Daniel Teta MD, PhD^j, Renuka Visvanathan MBBS, PhD^k, Elena Volpi MD, PhD^l, Yves Boirie MD, PhD^m



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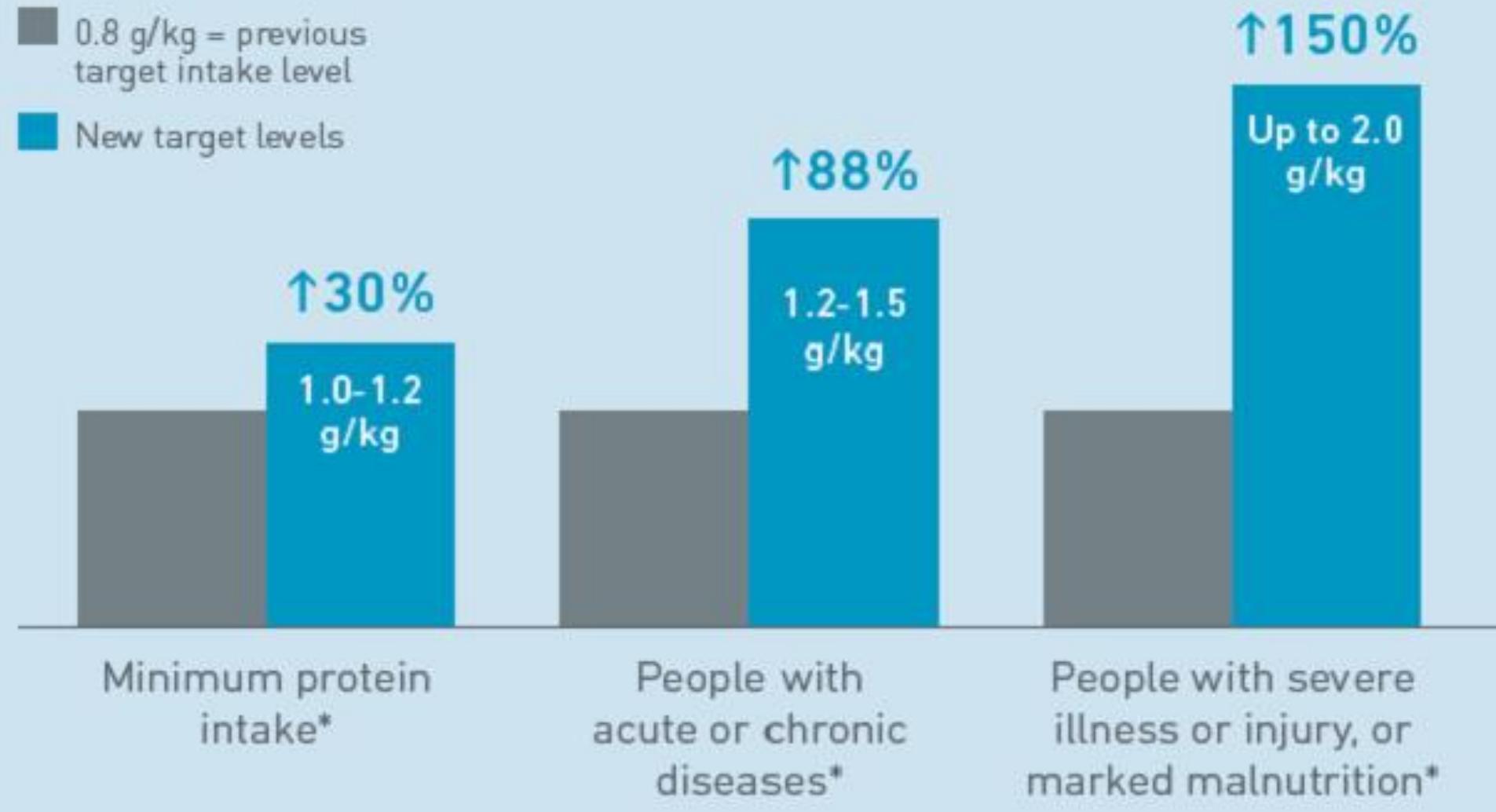
ESPEN endorsed recommendation

Protein intake and exercise for optimal muscle function with aging: Recommendations from the ESPEN Expert Group

Nicolaas E.P. Deutz^{a,*}, Jürgen M. Bauer^b, Rocco Barazzoni^c, Gianni Biolo^c, Yves Boirie^d, Anja Bosy-Westphal^e, Tommy Cederholm^{f,g}, Alfonso Cruz-Jentoft^h, Zeljko Krznarićⁱ, K. Sreekumaran Nair^j, Pierre Singer^k, Daniel Teta^l, Kevin Tipton^m, Philip C. Calder^{n,o}

New recommendations call for higher protein intake (g per kg of bodyweight) in those aged >65 years¹

- 0.8 g/kg = previous target intake level
- New target levels



*Caution needed among those with severe kidney disease (i.e. estimated Glomerular Filtration Rate <30mL/min/1.73m²), calculating their needs differently.

The SPRINTT study has some answers



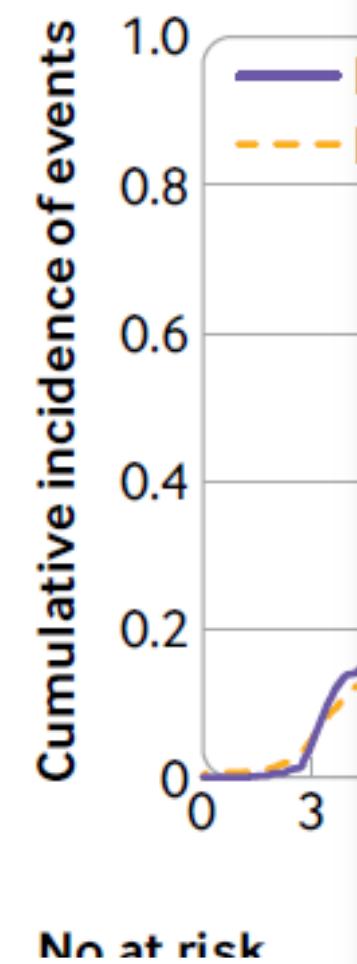
Innovative Medicines Initiative



- ECA, intervención multicomponente vs educación sobre envejecimiento sano, hasta 36 meses
- 16 centros en 11 países europeos
- 1519 hombre y mujeres en la comunidad con 70+ años, fragilidad física y sarcopenia (SPPB 3 a 9)
- Intervención multicomponente: ejercicio de intensidad moderada 2 veces a la semana en el centro y hasta 4 a la semana en casa + consejo nutricional personalizado
- Variable primaria: discapacidad para la movilidad (incapacidad de caminar independientemente 400 m en <15 minutos).



SPRINTT



Study	No with event/No in group (%)		Hazard ratio (95% CI)	Hazard ratio (95% CI)	Interaction P value
	Multicomponent intervention	Lifestyle education			
Overall	283/605 (46.8)	316/600 (52.7)	0.78 (0.67 to 0.92)		
Sex					
Men	88/171 (51.5)	97/175 (55.4)	0.87 (0.65 to 1.17)	0.38	
Women	195/434 (44.9)	219/425 (51.5)	0.75 (0.61 to 0.91)		
Race or ethnicity					
White	253/535 (47.3)	290/526 (55.1)	0.75 (0.63 to 0.90)	0.30	
Other	1/7 (14.3)	4/8 (50.0)	0.23 (0.03 to 2.15)		
Age (years)					
<80	115/318 (36.2)	154/336 (45.8)	0.75 (0.59 to 0.96)	0.94	
≥80	168/287 (58.5)	162/264 (61.4)	0.76 (0.61 to 0.95)		
History of CVD					
No	54/162 (33.3)	80/177 (45.2)	0.62 (0.44 to 0.89)	0.15	
Yes	229/443 (51.7)	236/423 (55.8)	0.84 (0.69 to 1.01)		
History of diabetes					
No	216/474 (45.6)	236/461 (51.2)	0.82 (0.68 to 0.99)	0.40	
Yes	67/131 (51.1)	80/139 (57.6)	0.70 (0.50 to 0.97)		
Gait speed (m/s)					
<0.8	226/450 (50.2)	248/444 (55.9)	0.79 (0.66 to 0.95)	0.96	
≥0.8	57/155 (36.8)	68/156 (43.6)	0.78 (0.54 to 1.12)		

Fig 3 | Prespecified subgroup analyses in participants with baseline short physical performance battery (SPPB) score of 3-7. CVD=cardiovascular disease; CI=confidence interval

Fármacos para la sarcopenia

Medications for the treatment of sarcopenia : a review

Context

Sarcopenia

= loss of muscle mass
and function



Widely prevalent and severe
condition in older adults



Recognized as a disease
ICD -10-CM code M62.84

No drug currently approved

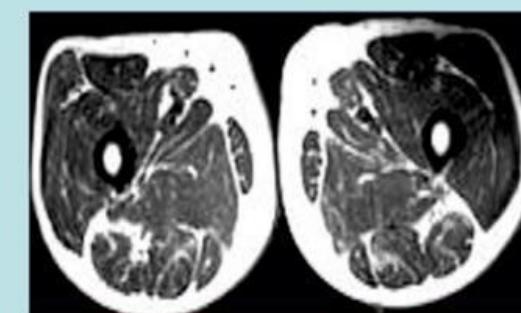
Yves Rolland et al. 2023
Metabolism, clinical and experimental

Results

Very few drug trials have targeted sarcopenic patients

Most drug trials were performed in elderly population
with varying characteristics

Efficacy was tested separately
on the clinical components of sarcopenia :



muscle mass



muscle strength



physical performances

- 
- 
- Testosterone
 - Selective Androgen Receptor Modulators (SARMs)
 - Estrogen
 - Dehydroepiandrosterone (DHEA)
 - Insulin-like Growth Factor-1 (IGF-1)
 - Growth Hormone (GH)
 - GH Secretagogue (GHS)
 - drug targeting Myostatin/Activin receptor pathway
 - Vitamin D
 - Angiotensin Converting Enzyme inhibitors (ACEi)
 - Angiotensin Receptor Blockers (ARBs)
 - β-blockers

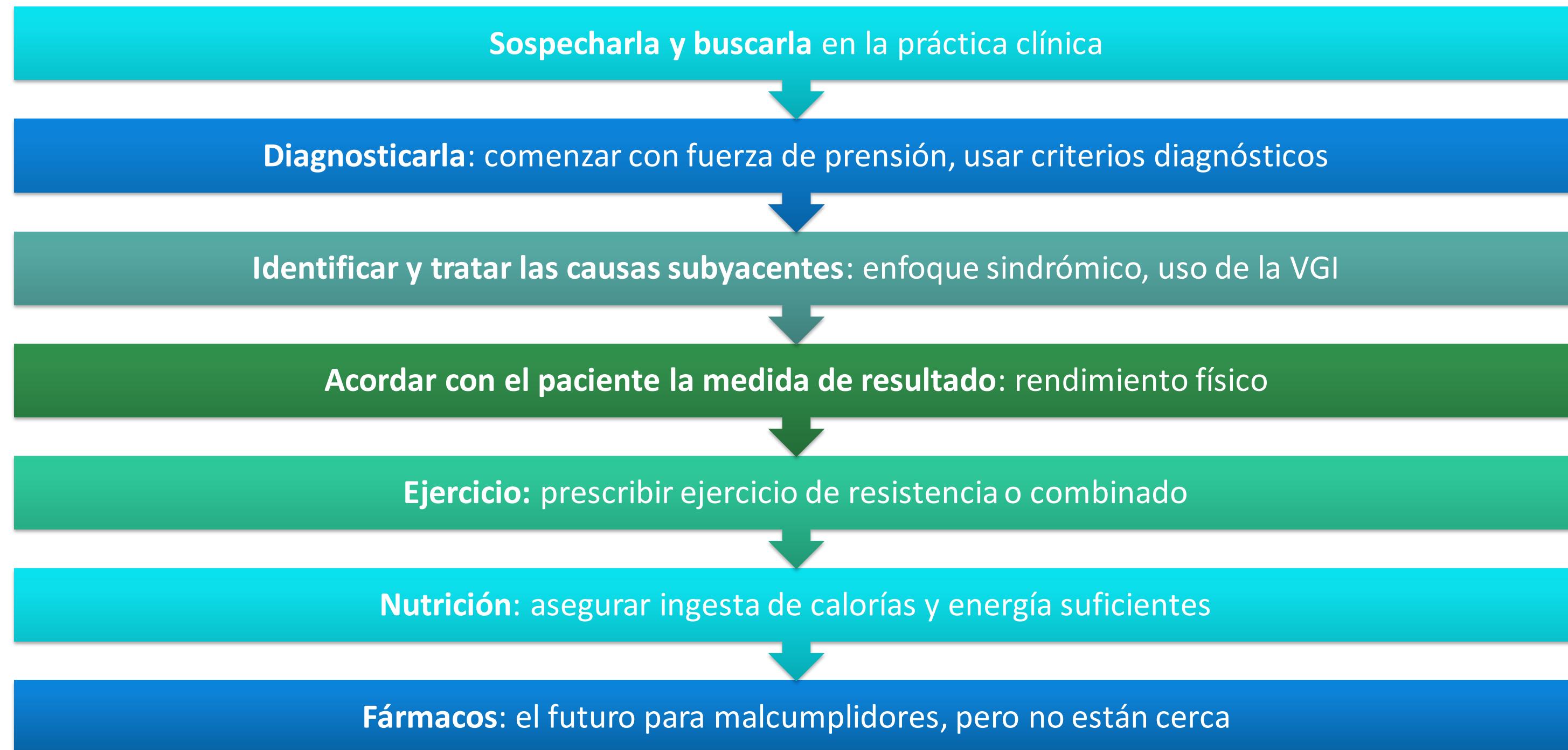
Conclusion

No drug translated into clinically
relevant improvements on
physical performance.



Diagnóstico y manejo de la sarcopenia

Mi enfoque personal (noviembre 2023)



GRACIAS