

# An overview of sarcopenia: facts and numbers on prevalence and clinical impact

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**Abstract** Human muscle undergoes constant changes. After about age 50, muscle mass decreases at an annual rate of 1–2%. Muscle strength declines by 1.5% between ages 50 and 60 and by 3% thereafter. The reasons for these changes include denervation of motor units and a net conversion of fast type II muscle fibers into slow type I fibers with resulting loss in muscle power necessary for activities of daily living. In addition, lipids are deposited in the muscle, but these changes do not usually lead to a loss in body weight. Once muscle mass in elderly subjects falls below 2 standard deviations of the mean of a young control cohort and the gait speed falls below 0.8 m/s, a clinical diagnosis of sarcopenia can be reached. Assessment of muscle strength using tests such as the short physical performance battery test, the timed get-up-and-go test, or the stair climb power test may also be helpful in establishing the diagnosis. Sarcopenia is one of the four main

reasons for loss of muscle mass. On average, it is estimated that 5–13% of elderly people aged 60–70 years are affected by sarcopenia. The numbers increase to 11–50% for those aged 80 or above. Sarcopenia may lead to frailty, but not all patients with sarcopenia are frail—sarcopenia is about twice as common as frailty. Several studies have shown that the risk of falls is significantly elevated in subjects with reduced muscle strength. Treatment of sarcopenia remains challenging, but promising results have been obtained using progressive resistance training, testosterone, estrogens, growth hormone, vitamin D, and angiotensin-converting enzyme inhibitors. Interesting nutritional interventions include high-caloric nutritional supplements and essential amino acids that support muscle fiber synthesis.

**Keywords** Sarcopenia · Muscle mass · Prevalence · Morbidity

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## 1 A history of muscle loss

Muscle is key to motion. As we age, significant changes in muscle mass and quality take place. After about age 50, muscle mass decreases at an annual rate of 1–2% [1, 2]. The decline in muscle strength is even higher, amounting to 1.5% per year between ages 50 and 60 and 3% per year thereafter. These findings have only received systematic research over the last two or three decades during which interest in age-related changes in physiologic function increased. This fact appears indeed curious because Macdonald Critchley, then junior neurologist at King's College Hospital in London, wrote already in 1931 that “the entire musculature tends with advancing age to undergo involutional changes, which are manifested as wasting” [3]. He went on saying that “probably the chief

cause of this change is to be demonstrated in the general process of senile atrophy, which shows itself in the muscles and elsewhere.” Later on, in the 1970s, Nathan Shock published a series of articles on age-related physiologic functions using data from the first large-scale longitudinal study in this field [4]. Altogether, it evolved that no decline in structure and function is more dramatic than the decline in muscle mass that develops as we age. Irwin Rosenberg realized that if this phenomenon were to be taken seriously, a name was required, and at a meeting in Albuquerque, New Mexico, in 1988, he suggested to use the term sarcopenia [5]. Following the recommendation by Morley, the term took hold over these last 20 years [6].

## 2 How to measure muscle mass and muscle strength?

The name sarcopenia is derived from Greek *sarx* (flesh) and *penia* (loss), literally meaning poverty of flesh. Sarcopenia is one of the four main reasons for loss of muscle mass, the others being anorexia, dehydration, and cachexia [7, 8]. It is difficult to estimate the prevalence of sarcopenia (Table 1), mostly because of practical difficulties in assessing muscle mass. Many different methodologies have been used over the last 20 years, and new techniques are still being introduced. On average, it is estimated that 5–13% of elderly people aged 60–70 years are affected by sarcopenia, and the numbers increase to 11–50% for those aged 80 or

above. Sarcopenia may lead to frailty, but not all patients with sarcopenia are frail. In essence, sarcopenia is about twice as common as frailty [9].

The broadness in the range of sarcopenia prevalence is partly due to the heterogeneity of study populations, but also due to the different techniques used to assess muscle mass. Dual-energy X-ray absorptiometry (DEXA) is currently considered the gold standard. The name is derived from the fact that two X-ray beams are used with different energy levels of minimal intensity [10]. Other methods used to measure muscle mass include bioelectrical impedance, computed tomography, magnetic resonance imaging, urinary excretion of creatinine, anthropometric assessments, and neutron activation assessments [7]. Depending on the actual technique used in different studies and on the cutoff values chosen, the prevalence of muscle mass may vary considerably (Table 1). Many institutions use handgrip strength as a standard measure for assessing muscle strength. Physical performance can be analyzed using simple and easy-to-do tests such as the short physical performance battery test [11], usual gait speed [12], the timed get-up-and-go test [13], or the stair climb power test [14].

## 3 Pathophysiological changes in sarcopenia

Using such techniques, it became clear that aging is associated with changes not only in muscle mass but also

**Table 1** Large-scale studies into the prevalence of sarcopenia

Cohort (country)	n (% female)	Age	Sarcopenia definition (assessment method)	Sarcopenia prevalence	Reference
CHS (USA)	5036 (56.4%)	>65 years	Categories of skeletal mass index, defined as muscle mass normalized for height (BIA)	Moderate sarcopenia, m: 70.7%, f: 41.9%; severe sarcopenia, m: 17.1%, f: 10.7%	[35]
EPIDOS (France)	1458 (100%)	All >70 years; mean 80.3±3.8 years	Appendicular skeletal muscle mass <2 SD below the mean of a young female reference group (DEXA)	9.5%	[36]
InCHIANTI (Italy)	1030 (54.5)	Range 20–102	Calf muscle cross sectional area more than 2 SD below population mean (CT scan)	m: 20% at 65 years, 70% at 85 years f: 5% at 65 years, 15% at 85 years	[37]
NHANES III (USA)	14818	>18 years; 30% >60 years	Skeletal mass index was defined as muscle mass/body mass x 100; sarcopenia class I defined as skeletal muscle mass 1–2 SD, sarcopenia class II defined as skeletal muscle mass >2 SD from the mean of young subjects (BIA)	In subjects aged >60 years: sarcopenia class I, m: 45%, f: 59%; sarcopenia class II: m: 7%, f: 10%	[21]
NMEHS (USA)	808 (47.3%)	m: 73.6±5.8 years; f: 73.7±6.1 years	Appendicular skeletal muscle mass <2 SD below the mean of a young reference population (substudy of DEXA)	<70 years, m: 13.5–16.9%, f: 23.1–24.1%; 70–74 years, m: 18.3–19.8%, f: 33.3–35.1%; 75–80 years, m: 26.7–36.4%, f: 35.3–35.9%; >80 years, m: 52.6–57.6%, f: 43.2–60.0%	[19]

BIA bioelectrical impedance assessment, CHS Cardiovascular Health Study, CT computed tomography, DEXA dual-energy X-ray absorptiometry, EPIDOS European Patient Information and Documentation Systems, NHANES National Health and Nutrition Examination Survey, NMEHS New Mexico Elder Health Study, SD standard deviation

in muscle composition, contractile, and material properties of muscle as well as in the function of tendons [15]. In aging muscle, there is a loss of motor units via denervation. These denervated motor units are recruited by surviving motor units, which puts an increased burden of work on them.

Altogether, there is a net conversion of fast type II muscle fibers into slow type I fibers with resulting loss in muscle power necessary for activities of daily living such as rising from a chair or climbing steps [15]. Other aspects include the deposition of lipids within muscle fibers. These effects—in contrast to cachexia [8]—do not lead to a net loss in body weight, but to a significant reduction in muscle strength. Indeed, in healthy volunteers, the maximal velocity during cycle ergometry decreased by 18% from the third age decade (20–29 years) to the sixth (50–59 years) [16]. Another 20% were lost between the seventh (60–69 years) and the ninth age decade (80–89 years) [16]. The loss of maximal oxygen consumption (peak  $\dot{V}O_2$ ) with increasing age has also been attributed to reduced muscle mass and cardiac output [15]. The risk of falls and consequently of fractures is closely related to reduced muscle mass as well: After multivariable adjustment, a 2.3-fold increase in the risk of falls was reported for patients in the lower third of handgrip strength as compared to the upper third [17]. Likewise, another large-scale study in more than 2,100 elderly subjects found that a low walking speed is an independent risk factor of falls [18].

#### 4 Making a diagnosis of sarcopenia

Using the above knowledge, it does not come as a surprise that the likelihood of having a physical disability is higher in elderly subjects who present with height-adjusted appendicular muscle mass 2 standard deviations below the mean of young adult as compared to those with normal muscle mass [19]. Having said that, the difficulty in making a correct diagnosis of sarcopenia is easily understood. Several definitions and diagnostic criteria have been proposed over the last 20 years. Many of them are not easily applicable in daily clinical practice.

A consensus definition formulated by experts from a vast array of different medical fields recently suggested to diagnose sarcopenia when two criteria are fulfilled: (1) a low muscle mass and (2) a low gait speed [20]. Normal muscle mass is defined using data derived from young subjects aged 18–39 years from the Third NHANES population [21], and the requirement for a diagnosis of sarcopenia is the presence of a muscle mass  $\geq 2$  standard deviations below the mean of this reference population. This value can normally be calculated automatically by

equipment such as DEXA-scanners. A low gait speed is defined as a walking speed below 0.8 m/s in the 4-m walking test [22]. The European Working Group on Sarcopenia in Older People suggested diagnosing sarcopenia when at least two of three criteria apply: (1) low muscle mass, (2) low muscle strength, and/or (3) low physical performance [23]. Cutoff points are defined in a similar manner as by the consensus group mentioned before, namely 2 standard deviations below the mean reference value for muscle mass and muscle strength of a reference population and a gait speed  $\leq 0.8$  m/s.

#### 5 Treatment approaches to sarcopenia

A diagnosis of sarcopenia remains a rare case. But even if the diagnosis is reached, the treatment of sarcopenia remains challenging. Many different approaches have been pursued, but exercise and physical activity are important considerations for both sarcopenia prophylaxis [24, 25] and sarcopenia management [26]. Progressive resistance training, performed 2–3 times per week by older people, has been shown to improve gait speed, timed get-up-and-go, climbing stairs, and overall muscle strength [27]. Nutritional interventions also have an important impact. Current recommendations state that protein should be consumed at a rate of 0.8 g/kg/day, but about 40% of persons above the age of 70 years do meet this amount [28]. Additional calorie intake of 360 cal per day together with resistance exercise training has been shown to increase leg muscle strength in nursing home residents after 10 weeks. Similar effects were described in cachectic patients [29]. Supplementation of essential amino acids has been shown to improve handgrip strength and 6-min walking distance in elderly subjects after 3 months [30]. Other therapeutic approaches include the use of testosterone, estrogens, growth hormones, vitamin D, and angiotensin-converting enzyme inhibitors [7, 25, 31]. In addition, animal studies have recently reported beneficial effects of soluble activin receptor type IIB (ActRIIB) treatment [32] and myostatin inhibition [33].

The first step in the sarcopenia journey is to create more awareness of this clinical problem, both by the general public and by healthcare professionals. In this respect, the implementation of standardized diagnostic criteria was extremely important. However, sarcopenia is a phenomenon that is present not only in healthy elderly subjects but also in those with chronic illnesses, such as chronic heart or renal failure. A debate exists as to whether muscle loss should be termed sarcopenia under these conditions as well. More prospective studies are required to understand the prevalence, incidence, phenotype, and the clinical impact of sarcopenia. Such studies are only

beginning to emerge. Furthermore, the discussion continues whether sarcopenia, i.e., loss of muscle mass, should be separated from dynopenia, i.e., loss of power. Also, as enthusiasts try to make sarcopenia all encompassing by adding a functional definition, the definition lines between frailty and sarcopenia are becoming blurred. There is a need for a consensus decision regarding the use of these terms.

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