Disparities in the consequences of sarcopenia: implications for African American Veterans

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INTRODUCTION

The phenomenon of age-related muscle wasting has been long recognized by geriatricians, rehabilitation specialists, and public health practitioners. However, Federal agencies are slow to recognize sarcopenia as a diagnostic term despite the ubiquitous observation of muscle loss as a signature event of aging. The formal adoption of sarcopenia as a clinical diagnosis has been complicated by its evolving operational definition and the varied approaches to the assessment of lean body mass (LBM). Currently, formal assessments of LBM rarely occur within the context of standard geriatric care. The failure to adequately assess muscle tissue within the larger context of geriatric medical care has significant public health implications given the importance of adequate muscle function for "successful aging" and the maintenance of independent living (Visser and Schaap, 2011).

The prevalence of sarcopenia is approximately 25% of the U.S. population over 70 years of age (Baumgartner et al., 1998; Lorenzo, 2009). Additionally, U.S. health care expenditures ascribed to sarcopenia, based on the Third National Health and Nutritional Examination Survey (NHANES III) data, have been conservatively estimated at 11.8 billion dollars (Janssen et al., 2004a). Given the national trends in the Veteran population with respect to age, reported disability levels, and increased representation of African American soldiers, the potential impact of sarcopenia on the Veterans Health Administration (VHA) healthcare system merits consideration.

AFRICAN AMERICANS AND THE VHA HEALTHCARE SYSTEM

The total Veteran population was estimated to be over 22 million in 2012 which also includes approximately 2.2 million women. There was significant minority representation among Veterans during this same period with African Americans members totaling 2.6 million men and 430,000 women. The 2011 Veteran Population Projection Model (VA Office of the Actuary, 2011) projects a nearly 35% decrease in the total Veteran population between 2013 and 2040. Nonetheless, this population trend will be countered by an anticipated concomitant increase in the percentage of minority Veterans from 21 to 34% during this same time period. African Americans will constitute the largest proportion of this projected demographic shift in the Veteran population (National Center for Veterans Analysis and Statistics, 2011). These changes reflect the advent of the All-Voluntary Force and recent military conflicts such as the Gulf War I and II which were notable for the high proportion of participating minority service members with nearly one in three soldiers being African American (Census Bureau, 2010). It is important to note that the

Veteran population is older than the general population with a median age of 62 in comparison with a non-Veteran median age of 43. The oldest Veterans reflect a larger proportion of Caucasians (Census Bureau, 2010; National Center for Veterans Analysis and Statistics, 2011) as their median age is 64 in comparison to minority Veterans who have a median age of 54-57. However, the relatively younger subset of minority Veterans coupled with the projected demographic changes suggest that the need to effectively manage geriatric syndromes in African Americans within the VHA healthcare system will continue to increase over time.

SARCOPENIA: DIAGNOSTIC CONSIDERATIONS FOR AFRICAN AMERICANS

Sarcopenia is an age-related syndrome characterized by a decrease in muscle mass and associated with a loss of strength and power, diminished functional performance, and increased disability (Newman et al., 2003; Morley et al., 2011). The observation that basic mobility and independent living status may be more dependent on lower extremity muscle performance rather than LBM in older adults (Visser et al., 2000; Kamel, 2003) was the impetus for some investigators to ascribe specific terms (e.g., dynapenia and kratopenia), to characterize the age-related loss of muscle force and power (Morley et al., 2011). Moreover, the Foundation for the National

Institutes of Health Sarcopenia Project has proposed to qualify sarcopenia as a form of skeletal muscle function deficit to reflect the varied etiology of mobility impairments (Correa-de-Araujo and Hadley, 2014). The recognition of total body mass or body fat as independent factors that influence functional status gave rise to sarcopenic index measures that combine these variables with estimates of LBM (Newman et al., 2003). Consensus groups such as the European Working Group on Sarcopenia in Older People (EWGSOP) (Cruz-Jentoft et al., 2010) have proposed a staged diagnostic criteria for sarcopenia that include elements of LBM, strength, and functional performance in response to these contemporary research findings (Table 1). These consensus groups have recommended customary walking speed as a key component of the sarcopenia screening criteria, and the Society for Sarcopenia, Cachexia and Wasting Disorders (SSCWD) has recommended that only older adults with

mobility deficits and/or a history of falls should be considered for sarcopenia screening (Morley et al., 2011). This screening model has potential shortcomings that may be cause for concern. Just as bone fracture is the signal event of osteoporosis, an observed functional limitation may be deemed a signal event of sarcopenia. Moreover, it remains an open question if these aforementioned conditions for sarcopenia screening, derived largely from population-based studies involving relatively healthy community-dwelling adults (Dam et al., 2014), are appropriate for usage among a Veteran population with a high prevalence of comorbid conditions.

RACIAL/ETHNIC CONSIDERATIONS FOR BODY COMPOSITION ASSESSMENT

Despite the recent evolution of the diagnostic criteria for sarcopenia, the assessment of LBM remains vital to our understanding and clinical management

of age-related muscle dysfunction. A critical issue to consider in addressing possible sarcopenia disparities in African Americans is the influence of racial/ethnic characteristics on common methods of body composition assessment. African Americans have approximately 5-8% higher levels of muscle mass in comparison to Caucasians based on total body potassium assessment (TBK), and this difference applies to both men and women and persists from childhood to old age (Ortiz et al., 1992). African Americans may also have 7-10% greater levels of bone mineral density (BMD) as measured with dual-energy X-ray absorptiometry (DXA), even when controlling for LBM and stature in men (Ettinger et al., 1997). However, the rate of age-related LBM and BMD loss may be similar between the two races (Cohn et al., 1977). In addition, the distribution of body fat differs between these groups as Caucasians have greater skinfold thickness in their chest, abdomen,

Table 1 | Categories of the sarcopenia syndrome based on causative factors, body composition, and stages of severity.

| Causative factors | | Body composition | | Staging | |
|---|--|--|---|----------------------|---|
| Primary Sarcopenia | Age-related only | Sarcopenia | LBM loss criterion only ^b | Pre- sarcopenia | LBM loss criterion only ^b |
| Secondary Sarcopenia Contributing comorbid factors and behavioral conditions | Activity-related Disease-related Nutrition-related | Sarcopenia class: aLM/h ² • Class I (−1 to −2 SD) • Class II (↓ −2 SD) | A range of criteria based on LBM (relative to stature) ^c | Sarcopenia | LBM loss criterion, and strength loss or functional status criteria met ^f |
| Myopenia All-cause designation, independent of age | LBM loss, affecting functional status or mortality | Sarcopenia class: SMI • Class I (−1 to −2 SD) • Class II (↓ −2 SD) | A range of criteria based on ratio of LBM to body mass (relative to body mass) ^d | Severe Sarcopenia | All criteria are met |
| Skeletal Muscle Function Deficit All-cause designation, independent of age | Diminished muscle performance ^a , affecting functional status | Sarcopenic obesity | LBM maintenance or loss, and body fat criterion met ^e | | |

^a Muscle performance as measured by a given measure or estimate of strength, power, or capacity (e.g., endurance/fatigue; Correa-de-Araujo and Hadley, 2014). ^b Appendicular lean mass (aLM/h²; men and women, respectively): 7.26 and 5.45 kg/m² (Baumgartner et al., 1998); 6.12 and 5.29 kg/m² (African American cohort; Kelly et al., 2009); aLM scaled to body mass index (BMI): 0.789 and 0.512 (aLM_{BMI}; men and women, respectively; Dam et al., 2014); use of a single criterion for sarcopenia involving lean body mass (LBM) may include adjustments for height, body fat, and/or body mass by some investigators (Cruz-Jentoft and Morley, 2012).

^c Class I: 8.51–10.75 kg/m² in men, and 5.76–6.75 kg/m² in women; Class II: ≤8.50 kg/m² in men, and ≤5.75 kg/m² in women (Janssen et al., 2004a,b). ^d Class I: 31–37% in men, and 22–28% in women; Class II: ≤30% in men, and ≤21% in women [SMI, skeletal muscle mass index; (muscle mass/body mass) * 100; Janssen et al., 2002].

^eBody fat percentage (BF%): 27% in men and 38% in women (Baumgartner, 2000); BMI: ≥30 kg/m² (Schrager et al., 2007); cohort specific criteria: 2 highest quintiles of BF% and 2 lowest quintiles of lean body mass (Gomez-Cabello et al., 2011).

^f Grip strength: <30 kg in men, and <20 kg in women (Murphy et al., 2013); <26 kg in men, and <16 kg in women (Dam et al., 2014); gait speed: <0.8 m/s (Cruz-Jentoft et al., 2010); gait speed or distance: <1.0 m/s or <400 m during the 6-min walk test (Morley et al., 2011).

and thighs, whereas African Americans have greater subscapular skinfold thickness (Zillikens and Conway, 1990; Nindl et al., 1998). These phenotypic patterns may confound body composition assessment methods that do not account for these factors. Consequently, an alternative LBM density value of 1.13 g/cm³ has been proposed for African Americans instead of the 1.10 g/cm³ value typically used in body composition models (Schutte et al., 1984).

The most common whole body tissue composition methods used in sarcopenia research include anthropometry, bioelectrical impedance (BIA), and DXA. Anthropometry is often used in community-health settings and constitutes a broad assessment category that features extremity circumferential measurements, body mass index (BMI), and skin fold measurements. While circumferential measurements have been shown to be associated with diminished activities of daily living (ADL) (Baumgartner et al., 1998), their use as proxy body composition measures systematically underestimate LBM in African Americans based on comparative differences in LBM density, body fat distribution, and even relative body proportions (Wagner and Heyward, 2000). BMI has served as a valuable predictor of critical health outcomes (Kane et al., 2011), but also does not account for racial/ethnic differences in LBM density or body fat distribution and may overestimate the obesity rates in African Americans (Kleerekoper et al., 1994). Skin fold measurements have limitations similar to the aforementioned anthropometric methods, and have also been shown to underestimate LBM density in African Americans (Schutte et al., 1984). In addition, BIA has been used extensively to characterize body composition in population-based studies (Janssen et al., 2002, 2004b; Chien et al., 2008). The interpretation of body composition data involving African Americans and BIA measures may be limited due to assumptions concerning body fat distribution and relative body proportions. However, Wagner et al. (1997) have identified a validated predictive model appropriate for use in this population. Their use of the Segal adiposity-specific equations, with expanded categories per the Stolarczyk modification, yields fairly accurate body composition measures for African Americans, underestimating FFM by 1.8 kg. While lacking the portability of anthropometry and BIA, DXA estimates of muscle mass have a high degree of association with TBK and the imaging method serves as a direct measure of BMD (Wagner and Heyward, 2000; Cruz-Jentoft and Morley, 2012). Therefore, DXA may not be unduly influenced by racial/ethnic differences in LBM density and may serve as an ideal method to assess sarcopenia in studies with multiracial participants and for clinical practice in hospital settings (sarcopenic LBM criterion values, Table 1).

THE PREVALENCE AND CONSEQUENCES OF SARCOPENIA IN AFRICAN AMERICANS

Despite the substantial clinical and financial burden attributed to sarcopenia, this geriatric syndrome remains an underdiagnosed condition and is rarely subject to a systematic screening process as a part of customary medical care for older adults (Fielding et al., 2011). The sarcopenia syndrome diminishes functional independence and results in a 3-4 times increased likelihood of developing a disability. In addition, sarcopenia may exacerbate hip fracture incidence due to the loss of requisite strength to maintain dynamic balance (Kamel, 2003). However, while some investigators have shown a significant relationship among diminished LBM, impaired lower extremity strength, functional performance, and disability (Malmstrom et al., 2013), others have reported that a negligible association exists between LBM and disability in community dwelling older adults (Visser et al., 2000). Walking speed, fall risk, negotiating stairs, basic mobility, and independent living status may be more dependent on lower extremity strength and power rather than LBM in older adults (Visser et al., 2000; Kamel, 2003).

The discordance between the agerelated loss of muscle mass and diminished function has been observed across ethnic groups. However, this non-intuitive clinical observation is most evident in African Americans. Few studies have reported epidemiological data by ethnic or racial group for sarcopenia. Investigators using data from the Health Aging and Body

Composition Study found that 14.3 and 18.4% of African American men and women, respectively, were classified as sarcopenic when adjusting for height and body fat (Newman et al., 2003). Others have noted a prevalence in Caucasians above 25% for men and women across all ages, whereas these values were reported as low as 6-12% in African Americans (Kan, 2009). Nevertheless, studies involving older, community-dwelling, African Americans consistently demonstrate that they have diminished functional performance, and higher levels of dependence with ADLs and risk of disabling conditions, in comparison to other racial/ethnic groups (Miller et al., 2005). What accounts for older African Americans having a higher prevalence of functional limitations and ADL difficulty despite their lower prevalence of sarcopenia?

Both Janssen and Visser have recognized high body fat levels as a confounder in the investigations into the association of LBM with functional limitations and disability (Janssen et al., 2002; Visser et al., 2002). The consideration of body fat and body mass adds an important dimension to the sarcopenia syndrome given that unchanging muscle mass coupled with increased body fat may portend significant functional and metabolic decline (Cruz-Jentoft and Morley, 2012). Moreover, myosteatosis (an increased proportion of intramuscular adipose tissue) has been observed in older adults of African ancestry, and this deleterious adaptation to aging may occur to a greater degree in comparison to Caucasians even when subcutaneous and visceral fat levels are similar among the groups (Song et al., 2004; Miljkovic-Gacic et al., 2008). While the higher risk of disabling conditions observed in African Americans likely involves an array of socioeconomic factors (Fuller-Thomson et al., 2009), high levels of myosteatosis are strongly associated with impaired lower extremity function and may serve as a possible explanation for the departure in linearity between muscle wasting and physical performance (Goodpaster et al., 2001).

Non-contractile features of muscle also affect health outcomes as age-related changes in skeletal muscle may be a critical progenitor of metabolic abnormalities (Albu et al., 2005). While preliminary

findings suggest that relative muscle mass may be more strongly associated with insulin resistance than strength (Bijlsma et al., 2013), there are divergent views on this point (Kamel, 2003; Sayer et al., 2005). Data from the NHANES III cohort reveal that sarcopenia is strongly associated with the homeostasis model assessment of insulin resistance and higher HbA1C levels in both non-obese and obese individuals under 60 years of age (Srikanthan et al., 2010). However, the relationship between sarcopenia and a higher prevalence of type 2 diabetes was only observed in obese individuals. Milikovic et al. (2009) determined that Afro-Caribbean men had higher myosteatosis levels and a lower body fat percentage in comparison to Caucasian men. Myosteatosis was associated with type 2 diabetes in both groups, but 44% of the Afro-Caribbean men were diagnosed with the metabolic disorder in comparison to 13% of the Caucasian men. While the negative impact of low muscle mass on glucose homeostasis cannot be discounted (Bijlsma et al., 2013), excessive intramuscular adipose tissue and intramyocellular lipid content may disrupt insulin receptor activity, limit glucose transport, and potentially result in adipokine release from local concentrations of adipose tissue (Miljkovic et al., 2009). The case could be made to incorporate age-related changes in muscle quality into the broad view of the sarcopenia diagnosis. However, more prospective study will be needed to better understand the role of myosteatosis in the pathogenesis of diminished insulin sensitivity and diabetes. Given that African Americans are 1.6-1.7 times as likely to have diabetes in comparison to Caucasians within the VA and civilian population, it may be a compelling interest of the Federal health agencies, professional and scientific organizations, and other stakeholders to study the impact of age-related muscle changes and how they interact with racial/ethnic group factors in health disparities (National Center for Veterans Analysis and Statistics, 2011).

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