



Published in final edited form as:

*Med Clin North Am.* 2018 January ; 102(1): 65–85. doi:10.1016/j.mcna.2017.08.007.

## Addressing Obesity in Aging Patients

**John A. Batsis, MD and Alexandra B. Zagaria, BA**

Section of General Internal Medicine, Dartmouth-Hitchcock Medical Center, 1 Medical Center Drive, Lebanon, NH 03756

Geisel School of Medicine at Dartmouth and The Dartmouth Institute for Health Policy & Clinical Practice, 1 Medical Center Drive, Lebanon, NH 03756

### Synopsis

Obesity in older adults impacts not only morbidity and mortality, but importantly impacts quality of life and the risk of institutionalization. Weight loss interventions can effectively lead to improved physical function. Diet-alone interventions can detrimentally impact muscle and bone physiology and without interventions to affect these elements, can lead to adverse outcomes. Understanding social and nutritional issues facing older adults is of utmost importance to the primary care provider. The insufficient evidence related to pharmacotherapy is discussed as well as an overview of using physiologic rather than chronologic age is addressed for identifying suitable candidates for bariatric surgery.

### Keywords

obesity; older adult; weight loss; physical function; pharmacotherapy; bariatric surgery; review

## Epidemiology of Aging & Obesity

By the year 2030 in the United States, over 20% of the population will be over the age of 65 years<sup>1</sup> (Figure 1), up from 15% of the population today<sup>2</sup>. The fastest growing demographic are the ‘oldest old’, individuals aged over 85 years. Much of the demographic shift is due to the emergence of baby boomers, adults born mid-1946 to mid-1964, into older adulthood (aged 65 years). Improvements in medical care, chronic disease management and infection control over the past century has also led to increases in life expectancy<sup>3,4</sup>. Based on recent census data, life expectancy at age 65 is now 82.8 years in males and 85.3 years in females<sup>5</sup>. The demographic changes observed during the aging process leads to a trajectory of disability<sup>6</sup>, independent of other influencing comorbidities. For instance, individuals surviving into old age are at risk of functional impairments – inability to transfer, walk, dress, eat, toilet, and bathe<sup>7</sup> – which subsequently lead a loss of independence, impairment

---

Corresponding author: John A. Batsis, MD, FACP, AGSF, Associate Professor of Medicine and The Dartmouth Institute for Health Policy & Clinical Practice, Dartmouth-Hitchcock Medical Center, 1 Medical Center Drive, Lebanon, NH 03756, Telephone: (603) 653-9500; Facsimile: (603) 650-0915, john.batsis@gmail.com.

**Publisher's Disclaimer:** This is a PDF file of an unedited manuscript that has been accepted for publication. As a service to our customers we are providing this early version of the manuscript. The manuscript will undergo copyediting, typesetting, and review of the resulting proof before it is published in its final citable form. Please note that during the production process errors may be discovered which could affect the content, and all legal disclaimers that apply to the journal pertain.

in quality of life and institutionalization<sup>8–10</sup>. Individuals are exposed to a longer period of time in which they may develop comorbidities and be at risk for developing incident disability<sup>11</sup>.

The obesity epidemic is not unique to a middle-age or a pediatric population. The prevalence of obesity in older adults, classified using body mass index (BMI), continues to rise over time. Recent estimates from the National Health and Nutrition Examination Surveys demonstrate that adults over age 60 have obesity rates exceeding 37.5% in males and 39.4% in females<sup>12</sup> (Figure 2). These estimates have been replicated and are increasing in other developed countries as well, including the United Kingdom<sup>13</sup> and Canada<sup>14</sup>.

## Defining Obesity in Older Adults

Body composition changes with ageing. Throughout adulthood, a natural increase in body fat develops up to the 8<sup>th</sup> decade of life, after which there is a reduction<sup>15</sup>. Redistribution of fat from peripheral and subcutaneous sources to a central location leads to increased waist circumference and waist-hip ratio in older adults. Importantly, there is a natural loss of muscle mass and strength with aging, termed sarcopenia<sup>16</sup>. Sarcopenia can also be accelerated in other processes including deconditioning, immobility or other acute illnesses<sup>17–19</sup>. Muscle mass and strength is believed to be vitally important in the preservation of physical function and independence in this population<sup>20</sup>. Using standard adult classifications for weight status, such as BMI, may thus underestimate adiposity for a given individual.

Many epidemiological and intervention-based studies have used BMI as a surrogate for adiposity. Standard BMI categories are used ubiquitously in clinical practice and are based on the World Health Organization cut-points of underweight (BMI<18.5kg/m<sup>2</sup>), normal weight (18.5–24.9), overweight (25–29.9), and obese (30+). BMI is easy to use, cheap and can be measured using a simple stadiometer and a weight scale; however, BMI is a poor measure of adiposity in older adults. First, individuals lose height as they age. The Baltimore Longitudinal Study of Aging noted that both males and females lost height with age<sup>21</sup> which impacts the BMI's denominator, possibly leading to an overestimation of its overall value in this population. Second, while BMI is clearly valuable as a population-level measure, it has poor diagnostic accuracy for identifying older adults with obesity<sup>22,23,24</sup>. Using data from the 1999–2004 National Health and Nutrition Examination Survey, the sensitivity of BMI in accurately identifying adiposity was 32.9% in males and 38.5% in females. Third, as previously described, older adults tend to gain weight in central regions of their body. BMI fails to distinguish between peripheral and visceral obesity, an important consideration in individuals who are classified as having normal weight central obesity<sup>25</sup>. Based solely on BMI, this category of individuals would often not be a target for obesity therapy because they fall within the normal range (BMI of 18.5–24.9kg/m<sup>2</sup>). Individuals with central obesity presenting with a normal BMI may also be at risk for adverse cardiometabolic dysfunction including dyslipidemia, coronary disease, hypertension, and early mortality<sup>26–28</sup>. One large scale epidemiological study using 15,184 adults aged 18–90 years with normal BMI and central adiposity based on waist-to-hip ratio<sup>29</sup> found that individuals with normal weight central obesity had a higher risk of total and cardiovascular mortality (HR of 1.87

[1.53,2.29] in males and 1.48 [1.35,1.62] in females). These relationships have been observed in older adults as well, whereby women have higher short-term cardiovascular mortality than males, and males have higher long-term cardiovascular mortality than females<sup>27</sup>. Irrespective of sex, individuals with normal weight central obesity (waist-hip ratio or waist circumference) are at higher risk of long-term disability<sup>28</sup>. Identifying and evaluating individuals in clinical practice who otherwise may not be identified as high risk is needed. Such misclassification can be problematic from a population-based management standpoint. Lastly, BMI accounts for both fat-free (muscle) and fat mass, the former of which declines during the aging process. This can further lead to misclassification and potential underestimation of adiposity and risk.

Unfortunately, gold-standard methods for identifying adiposity with accuracy, including magnetic resonance imaging and computer tomography, are non-covered indications in clinical practice unless performed for other reasons. Dual-energy absorptiometry is more routinely available, but assessment of body composition is not covered by Medicare. Bioelectrical impedance can be a crude measure for body composition assessment in older adults and portable devices are available; however, the alterations in fluid balance in older adults<sup>30</sup>, along with a higher incidence of prosthetic implants<sup>31</sup> and implantable cardiovascular devices<sup>32</sup> makes this modality less favorable. As such, BMI combined with a marker of central adiposity may provide a cost-effective approach to improved diagnostic accuracy within a clinical practice.

Established BMI cut-points correspond to adverse disease processes, including mortality<sup>33</sup>. Multiple population-based cohort studies have examined the relationship between obesity and premature death. In one study, obesity led to an estimated 111,909 excess deaths<sup>34</sup>. While obesity in mid-life is associated with reduced life-expectancy, the duration of obesity has a considerable effect on long-term mortality<sup>35</sup>, disability<sup>36</sup> and nursing home placement<sup>37</sup> as evidenced in numerous epidemiological studies. The relationship between BMI and mortality in populations has been shown to be representative of a “U” shaped curve<sup>34</sup>; individuals classified as underweight and obese are at the extremes with higher risks of mortality. With age, the curve flattens and shifts to the right, indicating that standard BMI cut-points differ in older adults than in younger populations. A meta-analysis that evaluated 32 studies between 1990–2013 (including 197,940 adults aged 65 and over) also demonstrated a “U” shaped curve<sup>38</sup>. Yet, the lowest risk of mortality was observed in those with a BMI of  $\sim 27.5 \text{ kg/m}^2$ . In fact, the risk of death only increases at a BMI  $> 33.0$  (HR 1.08 [95% CI: 1.00,1.15]).

In select older adult populations, not all participants with elevated BMIs should be considered at high risk. An ‘obesity paradox’ exists whereby elevated body weight may be protective, mitigating death in select populations. For instance, a nursing home systematic review evaluated 19,538 subjects with a BMI  $30 \text{ kg/m}^2$  and found that the risk of death was significantly lower than for the referent (normal BMI) group (HR 0.69 [95% CI: 0.60,0.79];  $p < 0.001$ )<sup>39</sup>. A number of factors are thought to explain this: first, issues related to the inability of BMI to discern between visceral and subcutaneous fat; second, cardiovascular fitness likely moderates the relationship between obesity and death. Individuals with high levels of cardiovascular fitness irrespective of their obesity

classification portend to better outcomes<sup>40</sup>; third, excess adiposity in high risk populations at risk for frailty, itself a predictor of death, may be protective. Populations such as hemodialysis patients, congestive heart failure, or nursing home residents all tend to lose weight (consisting of both fat and muscle) with aging which promotes wasting, cachexia and mortality<sup>41</sup>. Fourth, there may be self-selection—individuals with excess adiposity who have survived to old age may have a survival advantage as compared to those who have died earlier in life. As such, practitioners should be made aware of these considerations in select populations, particularly when using BMI as a measure for adiposity in older adults with these specific co-morbidities.

## Impact of Obesity on Physical Function/Disability

Overweight and obesity predisposes to disability and decreased physical functioning. Using the Health, Aging and Body Composition data, adults classified as overweight or with obesity using BMI at ages 25, 50 and 70–79 years had a HR 2.38 of incident disability over a 7 year period<sup>35</sup>. Similar relationships have been observed using either waist circumference or body fat percent, both in males and females. A systematic review by Schaap et al<sup>42</sup> demonstrated that adults with a BMI  $\geq 30\text{kg/m}^2$  aged 65 years and older had a 60% higher risk of incident disability (95% CI: 1.43, 1.80). A “U” shaped relationship is also observed between BMI and nursing home admission from community-dwelling adults<sup>43</sup>. Longitudinally, there appears to be a relationship. As the obesity epidemic has emerged in the past few decades, recent evidence suggests that its relationship with disability continues to be problematic, yet may be leveling off<sup>44</sup>. A recent study evaluated three consecutive time periods using National Health and Nutrition Examination Surveys (1988–1994, 1999–2004, and 2005–2012) and evaluated the association between obesity and disability. The population attributable fraction for obesity having a functional impairment and severe ADL impairment was 23.2% (95% CI: 20.5, 25.7) and 24.6% (95% CI: 12.3, 35.2), respectively, in individuals aged 60 years and older. Individuals classified as having obesity were still at much higher risk of limitations, yet limited wave-to-wave variability was observed. Other measures of adiposity, including body fat percent and waist circumference have also been associated with impaired physical function and disability and parallel these estimates. A study evaluating the cross-sectional association between obesity using body fat and disability demonstrated a significant odds of disability<sup>45,46</sup>. While there are challenges in the diagnostic accuracy of obesity, irrespective of the body composition or anthropometric measure used, in community dwelling adults, obesity is associated with a poorer prognosis of physical function.

Obesity is also associated with increased risk of falling in older adults. Over one-third of adults aged 65 years and older fall each year<sup>47</sup>, making it necessary to screen for fall risk in the primary care setting<sup>48,49</sup>. Two studies using Health and Retirement Study 1998–2006<sup>50</sup>, and the Behavioral Risk Factor Surveillance System 2014 data<sup>51</sup>, demonstrated the degree of obesity (class I vs. II vs. III) was associated with a higher risk of falling. Using the Health and Retirement Study data, those with class III obesity ( $\text{BMI} > 35\text{kg/m}^2$ ) have a OR 1.50 [1.21–1.86] of falling, while estimates from the Behavioral Risk Factor Surveillance System are slightly lower (1.23 [1.13, 1.35] in females; 1.18 [1.06, 1.32] in males). While fall risk is increased, risk of hip fractures from obesity is lower in this population<sup>52</sup>. Evaluating

individuals who are at risk can prevent falls that may otherwise lead to restriction of social function, fractures and death<sup>48,50</sup>.

## Sarcopenic Obesity – a subset of high-risk individuals

Sarcopenia in individuals with obesity is a subgroup that deserves specific attention. Sarcopenia is derived from the Greek word, “sarcos” meaning flesh and “penia” meaning lack of. Infiltration of fat occurs within muscle tissue that can lead to impairments in muscle physiological parameters<sup>53,54</sup>. The definition of sarcopenia and obesity (sarcopenic obesity) continues to be fraught with methodological challenges<sup>55</sup> and discrepancies in defining sarcopenia (muscle mass vs. muscle strength) and obesity (body fat vs waist circumference vs BMI). Nonetheless, such individuals are at considerably higher risk than those with either of the two conditions independently. While the medical definition has evolved over the past three decades<sup>55</sup>, based on the 2014 Foundations for the National Institutes of Health Conference<sup>20</sup>, sarcopenia is simply the loss of muscle mass and function with aging. Specific cut-points have been developed for use. While several earlier studies focused on loss of muscle mass as the key determinant of sarcopenia<sup>56</sup>, emerging evidence suggests that muscle strength may be a more powerful determinant of incident disability<sup>57</sup>.

The development of sarcopenia in old age is a natural phenomenon that can be partially mitigated with lifestyle interventions altering the threshold at which disability ensues<sup>16,58</sup>. Short- and long-term changes of both muscle mass and strength occur. Yet, changes in strength may occur without corresponding changes in muscle mass as observed in an earlier study using Health, Aging, Body Composition data<sup>59</sup>. Earlier work using the New Mexico Aging study demonstrated that individuals with sarcopenic obesity had a HR 2.63 [95%CI: 1.19–5.85] of developing an impairment of their instrumental activities of daily living over the course of a 8 year period<sup>60</sup>. This study defined sarcopenic obesity using appendicular skeletal muscle mass and body fat cutpoints<sup>61</sup>. Data from the InChianti study demonstrated that individuals with low muscle strength and obesity (based on knee extensor strength and BMI) had lower walking speeds than their counterparts over a 6-year period of time<sup>62</sup>. Much of our own work has revolved around using dual x-ray absorptiometry-defined muscle mass with the FNIH cutpoints with body-fat defined obesity (men>25%, females>35%), suggesting a significant relationship with limitations<sup>63</sup> and mortality<sup>64</sup>. Mortality is less clear as demonstrated using NHANES III data<sup>65</sup> (using BIA-defined muscle mass) and NHANES 1999–2004 data (using dual x-ray absorptiometry-defined muscle mass)<sup>64</sup>. Impaired muscle strength in conjunction with obesity, irrespective of its definition, is associated in cross-sectional and longitudinal studies with adverse and negative outcomes, more so than muscle mass<sup>42</sup>. Thus, the importance of identifying patients with both sarcopenia and obesity is of paramount importance.

## Evidence for Weight Loss in Older Adults?

Previous epidemiologic studies have provided conflicting findings on outcomes following weight loss in older adults; however, these studies failed to differentiate between intentional vs. unintentional weight loss and do not account for important confounding variables and reverse causality<sup>66,67</sup>. A joint consensus statement, published in 2005 by members of The

Obesity Society, American Society of Nutrition and The National Association for the Study of Obesity, provided some evidence on managing this disease in older adults,<sup>68</sup> and several randomized controlled trials have since been published demonstrating the benefits and the harms of weight loss in older adults<sup>69</sup>. (Note that we define ‘weight loss’ as intentional in the section below.)

A healthy lifestyle has been shown to compress the number of years of disability, according to data from the Cardiovascular Health Study 1989–2015<sup>70</sup>. Monitoring multiple lifestyle factors, including physical activity, diet, and BMI, 5,248 community adults aged 65 years and older who were not wheelchair dependent were identified. Activities of Daily Living were assessed and the ratio of the number of years living without any disability to the total number of years lived was ascertained to indicate compression or expansion of the disabling period. Obesity was associated with a decrease of 7.3% (95% CI: 5.4–9.2) as compared to normal weight individuals. The lowest quintile of the Alternative Healthy Eating Index was associated with a 3.7% (95% CI: 1.6–5.9) lower score than the highest quintile. Yet, engaging in physical activity demonstrated that for every 25 blocks walked in a week, 0.5% (95% CI: 0.3–0.8) higher proportion of years was gained disability-free. The article concluded that healthy lifestyles can compress the duration of disability in a person’s remaining lifetime.

A recent qualitative systematic review evaluated six randomized controlled trials from 2005–2015 in adults aged 60 and older (mean age >65 years)<sup>69</sup>. Obesity was defined as BMI 30kg/m<sup>2</sup> or waist circumference 88cm/102cm in females/males respectively. Of 5,741 citations, 19 trials were identified, of which 6 were unique cohorts. Results suggested that a dietary weight reduction program combined with a physical activity program (aerobic and resistance) improved physical performance and quality of life and lowered the risk of reduced muscle mass, strength and bone loss. Weight loss ranged from 0.5 to 10.7kg. A recently published randomized trial with 141 participants, demonstrated that weight loss inclusive of both aerobic and resistance exercises led to improvements in the physical performance test, peak oxygen consumption, and muscle strength, despite only marginal reductions in lean muscle mass<sup>71</sup>. Additionally, there was no difference in exercise-related adverse events. In a separate review, weight loss in older adults with obesity was associated with reduced risk of death<sup>72</sup>. The authors identified 15 randomized controlled trials including 17,186 participants with a mean age of 52 years at randomization. The mean BMI was 30–46 kg/m<sup>2</sup> with an average follow-up period of 27 months (ranging from 18 months to 12.6 years). The weight loss group experienced a 15% lower all-cause mortality risk (relative risk 0.85 [95% CI:0.73,1.00]). While further evidence is needed to ascertain the impact on long-term mortality, weight loss appears safe and effective in older adults with obesity.

## Cautions of Losing Weight in Older Adults

There are important risks that often get overlooked in this population by practitioners. Loss of weight leads not only to loss of fat mass, but also to loss of muscle mass, thereby promoting sarcopenia and its ensuing adverse outcomes<sup>73</sup>. The general principle that each pound lost equates to 75% fat and 25% muscle has been debated but is generally accepted<sup>74</sup>.

Moreover, loss of weight impacts bone metabolism and turnover promoting osteoporosis<sup>75-77</sup>.

Although sarcopenia is a natural phenomenon of the aging process, its acceleration with weight loss efforts is of considerable concern. Diet or diet and exercise-induced weight loss induces hormonal changes that negatively impact muscle mass and strength and is exacerbated by moderate caloric restriction. A review of 33 interventions demonstrated significant decreases from baseline in knee extensor strength (-7.5%) and handgrip strength (-4.6%) following diet-induced weight loss with moderate energy restriction<sup>76</sup>. Failure to engage patients in isokinetic resistance exercises will lead to likely loss of muscle mass and strength and reduce the impact of the gains in function individuals may otherwise attain with weight loss<sup>78</sup>.

A number of randomized controlled trials have demonstrated that caloric restriction alone leads to loss of bone mineral density (BMD). Villareal's group demonstrated that a loss of hip BMD was attenuated, in part, when resistance exercise was coupled with a weight loss program, preventing an increase in bone turnover<sup>58,71</sup>. In another study, a meta-analysis of 32 randomized controlled trials, weight loss had no significant effect on total BMD<sup>75</sup>. However, the pooled study data suggested that hip and lumbar spine BMD were significantly lower after 4 months, particularly in adults who were classified as having obesity. Lumbar spine BMD was also lower after calorie restriction in interventions longer than 13 months. While these results were in adults age 50 years and older, their sensitivity analysis stratified by age revealed that hip BMD loss was highest in adults age 65 and older. This finding has considerable implications for the older adult who is at risk of falling. Approximately 30% of falls among older adults result in injury, 10% of which are fractures<sup>79</sup>. Hip fractures are especially devastating in this population and portend considerable morbidity and mortality<sup>80</sup>.

Generally, obesity in the older adult population is safe and effective and can lead to considerable improvements in cardiometabolic risk and physical function. While there are important known risks associated with weight-loss in this population, they likely can be mitigated with appropriate health promotion interventions. As with all patients in a geriatrics practice, practitioners need to manage the benefits vs. the risks of any interventions, and in select individuals, weight loss should be encouraged.

## Obesity in the Primary Care Setting

Obesity prevention efforts should be based in primary care settings, where front-line clinicians have longitudinal relationships to provide brief, motivational interviewing to engage patients in behavioral change. Intensive behavioral counseling can induce clinically meaningful weight loss of between 0.3–6.6kg, but little research is available on primary care practitioners providing this care. A systematic review suggested that different interventionists can deliver counseling, both in person or by telephone in this setting<sup>81</sup>. An additional review suggested that a multidisciplinary team approach consisting of collaborative care was much more effective<sup>82</sup>. This review conflicts with a recent two-arm randomized trial of 2,728 patients that demonstrated that a behaviorally informed, very brief

(30 second) physician-delivered opportunistic intervention was acceptable and effective to reduce population mean weight (1.43kg [95% CI: 0.89,1.97])<sup>83</sup>.

In 2011, in the United States, the Centers for Medicare and Medicaid Services began reimbursing obesity counseling (current procedural terminology code G0447) for clinicians in a primary care setting to provide 22 targeted, 15-minute intensive behavioral therapy counseling in a continuous 12-month period<sup>84</sup>. The goal was to achieve a mean weight loss of 3kg in beneficiaries whose BMI was  $\geq 30$  kg/m<sup>2</sup>. Practice management barriers<sup>85</sup> exist in implementing this benefits, although novel technologies<sup>86</sup> may be helpful in addressing these issues. In 2012, the first full year of data, 27,338 (0.1%) of Medicare beneficiaries over age 65 availed of the benefit<sup>87</sup>. This increased slightly in 2013 to 46,821 (0.17%). The estimated proportion of persons with obesity using the benefit increased from 0.35% to 0.60%, with a mean of 1.99 and 2.16 claims/user. This data suggests its low uptake may not only be due to poor implementation patterns, but other support staff such as health coaches and dietitians should be integrating in delivering this important service. Novel delivery modalities to engage patients may be a strong consideration moving forward that include transition to value-based care models or increased reimbursements.

## Medical Evaluation Specific to Older Adults with Obesity

The nutritional needs and caloric intake for healthy older adults is known to decrease with age in both sexes. The caloric difference between early adulthood and older adulthood ranges between 300–500 kCal/day. Much of this is due to age-related phenomena related to basal metabolic rate which drops considerably with age<sup>88</sup>. The following section will discuss specific concerns that a primary care provider should consider using a ‘geriatric specific’ approach, as compared to a middle aged adult with obesity<sup>89</sup>.

Communication in clinical settings with older adults requires careful communication strategies that are often overlooked by clinicians. Behavioral techniques must be adapted to accommodate not only the sensory deficits the older adult faces such as hearing and vision, but need to be adapted to changes in cognition and executive function<sup>90</sup>, and their preference for shared-decision making<sup>91</sup>. Others have noted of a ‘gap’ between intentions and behavioral change<sup>92</sup>, which itself can be predicted by measures of executive function<sup>93</sup>. An inability to implement intentions will lead to poor execution of the desired change<sup>94</sup>. Older adults may also be better focused on single health behavior change interventions as compared to multiple<sup>95</sup>, focusing on a specific content. These approaches are less confusing and can be understood much more adequately. On the contrary, though, complex conditions in older adults require multiple strategies to deliver health promotion efforts which can be challenging to the clinician. Engaging individuals in strategies to improve self-efficacy through social support and change can be helpful to provide information to set goals, engage in change, and to promote self-monitoring<sup>96</sup>. Researchers continue to caution clinicians in applying behavioral change principles to older adults as they may require adaptation from a younger population.

While primary care providers care for older adults, the lack of specific geriatric training can be problematic in delivering behavioral change interventions. First, motivational



interviewing, a core tenant in eliciting change and in the Medicare Obesity Benefit is heavily influenced by the contextual aspects of delivery and by the clinician<sup>97</sup> – internists may approach elements differently than geriatricians. Second, geropsychological principles are often not integrated in routine interventions<sup>98</sup>. This includes social participation which is strongly related to better health and can lead to forming new goals in one's life<sup>99</sup>. This allows for a reframing of the discussion to engaging in change. Third, goals in seniors are different, in part due to multimorbidity<sup>100</sup>, but also due to the changing perceptions on aging and health<sup>101–104</sup>. Fourth, aging individuals have limited lifespans and, hence, patients focus more on the present day than the future. These elements lead to significant challenges in a busy primary care setting.

A thorough medical evaluation is needed in older patients that wish to improve their health and physical function through weight loss. The primary care clinician (or obesity medicine specialist) should identify whether there are any recent changes in health status (medical or economic). Recent hospitalizations and changes in functional status (e.g. joint replacement leading to physical inactivity) that may lead to weight gain<sup>105</sup>. Other standard questioning on weight history, previous strategies and alcohol use parallel methods used in the general population. Notably, we highlight two main concerns specific to older adults including medications and social support.

The number of medications prescribed increases with age and with the number of chronic conditions. Polypharmacy is a significant risk in older adults as it is associated with increased risk of cognitive impairment, urinary incontinence, falls and declines in physical function<sup>106</sup>. The American Geriatric Society has developed the Beers Criteria<sup>107</sup>, which clearly identifies potentially inappropriate medications in older adults and assists healthcare providers in improving medication safety for this population. Its purpose is to inform clinical-decision-making concerning the prescribing of medications for older adults in order to improve safety and quality of care. There are a number of medications that are considered 'obesogenic' that should be re-evaluated as part of the evaluation. As older adults have a higher incidence of diabetes<sup>108</sup>, depression<sup>109</sup>, pain<sup>110</sup> and hypertension<sup>111</sup>, medications treating these chronic diseases may increase the risk of a person gaining weight. Additionally, a number of these medications are listed on the Beer's criteria. The primary care practitioner, in concert with a clinical pharmacist on the multidisciplinary team can assist in streamlining not only the number of medications, but also the class of medications that promote weight gain. Table 1 briefly highlights some of the commons medications that predispose to weight gain.

Socioeconomic and ethnic disparities are two specific social determinants of health that increasingly are being recognized as important predictors in obesity management and adverse health. An elder on a fixed income (often a social security income) and must make choices between their food consumption and other care needs. Food insecurity is defined as the limited or uncertain availability of nutritional adequate and safe foods, or the limited or uncertain ability to acquire acceptable foods in socially acceptable ways<sup>112</sup>. Food insecure older adults individuals have poorer dietary intake, nutritional status and health status than food secure older adults<sup>113</sup>. Simple questions that should be asked include:

- Where is your food coming from?
- Who purchases your food?
- Do you have to pay for your medical bills and scrimp on food?

Such questions are helpful in that it leads to information on affordability of food. A true multidisciplinary team led by the physician that embodies a dietician, social worker and care manager can be helpful in intervening in this population. A dietician is of utmost importance in that they are able to perform, not only the usual functions of evaluation and counseling, but can assist this population in engage in substituting choices that they can afford.

Most established trials have evaluated calorie-restricted diets ranging from 500–750 kCal/day under the guidance of a registered dietician<sup>69</sup>. In older adults, there is ample evidence that diets such as the Dietary Approaches to Stop Hypertension<sup>114</sup> or the Mediterranean diet<sup>115</sup> have demonstrated improvements in metabolic parameters, weight loss, long-term disability, mortality and cognition. Very low energy or protein sparing diets should be avoided in older adults due to risks of dramatic fluid and electrolyte shifts and proportionally can lead to augmented muscle mass loss. Villareal, in his studies, proposed a loss of ~10% of baseline weight at six months<sup>58,71</sup>. Supplemental vitamin D of ~800–1000units and 1200mg of calcium should be considered. The latter can be from dietary or supplement sources, although we emphasize the consumption from dietary calcium if at all possible considering the recent controversies. Protein intake should also be augmented in older adults. Recommended dietary allowance is 1.0g/kg/day, yet since older adults produce less protein than younger persons, this should be revised to 1.0–1.2g/kg/day<sup>116</sup>. Also, a larger amount of protein is required to produce an equal response<sup>116</sup>. Notably, a recent Cochrane review demonstrated that protein supplements did not lead to improved outcomes<sup>117</sup>. Early pilot studies during weight loss interventions suggest improved short performance physical battery measures. The MeasureUP<sup>118</sup> in 67 subjects demonstrated, at 6-months, a  $-8.7 \pm 7.4$ kg and  $-7.5 \pm 6.2$ kg weight loss in the protein and control groups, respectively. The authors observed improvements in physical function based on the short physical performance battery of  $+2.4 \pm 1.7$ units and  $+0.9 \pm 1.7$  units, respectively ( $p=0.02$ ). Future studies should determine the amount of protein intake, the type of protein (meat vs. plant-based) and whether advised supplements for the treatment of sarcopenia (whey, leucine carnitine) can augment physical function and further mitigate sarcopenia.

Primary care providers have little training in counseling on physical activity<sup>119</sup>. They can, however, provide targeted information on exercise prescriptions. Such prescriptions should be individually tailored based on the individual's functional status and capacity. The American College of Sports Medicine suggest at least 150 minutes of low to moderate intensity aerobic activity in all patients, including older adults<sup>120</sup>. Seniors who may not have sufficient cardiovascular fitness may subdivide this time period into smaller increments to not only build their endurance but assist in ongoing behavioral change. The practitioner and their team can encourage patients to slowly increase their capacity to do so. During weight loss efforts, resistance exercises are of paramount importance for the prevention of sarcopenia. All individuals should engage in flexibility, balance, and strengthening activities with resistance bands or weights that can be free or be attached to wrists or legs. In those

with financial difficulties, we advise individuals to utilize common household items (i.e. soup cans or jugs of milk). The weekly goals are 2–3 days, with at least 48 hours rest between sessions. Each of these sessions should last between 30–40minutes, rotating muscle groups and exercises, with 10–12 repetitions for each sessions. Fatigue normally occurs between 8–12 repetitions and a bit of fatigue is advisable, but it is important to prevent injuries by starting low and going slow, starting with 1–2 pounds or with the lowest resistance color band. Physical therapists can assess a person’s 1-repetition maximum and advise advancement. We advocate that current multidisciplinary physical activity programs should be based on the Life Study<sup>121</sup>, a multi-center randomized controlled study that demonstrated reduced disability over a mean 2.6 year follow-up in 818 patients. Compared to a self-instructed exercise program, participants in this structured program demonstrated a 28% reduction in incident major mobility disability (95% CI:0.57,0.91). These materials are freely available on <<https://www.thelifestudy.org>>. Additionally, the National Institute on Aging has a booklet of exercises that can be accessed and downloaded at <<https://www.nia.nih.gov/health/publication/exercise-physical-activity/introduction>>, free of charge for clinicians and patients to engage at home.

Clinicians can play a paramount role in the monitoring of their older patients undergoing weight loss. While there is no firm evidence pertaining to monitoring parameters, we recommend the following. First, consideration should be given to assess baseline bone density during such efforts. Medicare indications for women are quite broad (ie: post-menopausal) yet those for men may be slightly more challenging to find. The United States Preventive Services Task Force recommendations for osteoporosis screening in men do not provide a firm statement and conclude that the evidence is insufficient to assess the balance of benefits and harms of screening<sup>122</sup>. Some potential indications in men include: x-ray evidence of possible osteopenia, osteoporosis or vertebral fractures; a person taking steroids; hyperparathyroidism; monitoring if osteoporosis drug therapy is working. Some examples of biomarkers of bone turnover that can be considered in the evaluative process, including osteocalcin, type I procollagen, urine collagen type-1 cross-linked N-telopeptide. These may help direct the impact of weight loss on bone turnover. Baseline and longitudinal monitoring of grip strength is an office procedure that can be easily integrated in any busy practice. The recent approval of an ICD-10 code for sarcopenia may allow for further screening and routine integration in practice. Last, monitoring of vitamin D levels may be helpful. Different societies have different views on monitoring of levels<sup>49</sup>. Our practice is to assess baseline Vitamin D levels during routine weight loss management. As with any intervention in older adults, risks vs. benefits of monitoring is needed, and further studies should best inform the appropriate indices clinicians should considering during such efforts.

## Pharmacotherapy

With the emergence of newer medications effective in weight management, older adults are increasingly asking about the possibility of taking such medications. The AACE/ACE guidelines explicitly state that there is insufficient evidence to recommend weight-loss medications in older adults<sup>123</sup>. As is the case with most pharmaceutical-based clinical trials, to prove efficacy, older adults were excluded from most trials, biasing outcomes towards younger patients. Unfortunately, we caution extrapolating such results to older adults whose

pharmacokinetics and pharmacodynamics properties differ from those in younger, robust populations who often have fewer comorbid conditions and are on fewer medications.

Of the obesity medications available, two trials (phentermine/topiramate and liraglutide) have documented efficacy analyses between older adults and younger counterparts. These trials enrolled 7% (n=254) and 6.9% (n=232) older adults, respectively, of their study subjects<sup>123–128</sup>. There were no observed differences in efficacy, safety and pharmacokinetics between subgroups. The other commercial medications had insufficient study sample to make any statistical comparisons. Preliminary data suggest that pharmacokinetic data on phentermine/topiramate, liraglutide, lorcaserin or naltrexone/bupropion were no different between younger/older. The propensity to add to an older adult's polypharmacy should not be overstated as this leads to medication errors and subsequent adverse drug events, increased risk for falls, delirium and costs. Table 2 highlights some of the absolute contraindications to these medications and some of the common side effects and major risks that older adults may face, irrespective of the lack of evidence. Given the need for obesity treatments and the limited data currently available, further research in older adult populations is important.

## Bariatric Surgery

An effective treatment approved by the NIH in 1991 is bariatric surgery<sup>129</sup>. This procedure has gained considerable popularity and is increasingly being performed in persons with obesity who are at high risk of medical complications and/or have co-morbidities. In the general population, there is considerable epidemiologic and trial data demonstrating its safety, efficacy, and effectiveness<sup>130–135</sup>. The extent of the safety and efficacy in older adults continues to be debated in the surgical literature. Many studies have used varying cut-points for older adults (ranging from 50–65 years), and are fraught with considerable methodological problems, including reduced study power, study time period bias, and inconsistent definitions. Additionally, the evolving surgical and medical care of this patient population, and the establishment of high volume Bariatric Surgery Centers of Excellence have led to considerably improved outcomes<sup>136,137</sup>. A number of systematic reviews have been published discussing the short- and long-term outcomes of bariatric surgery in an older adult population<sup>138</sup>, and are outside the scope of this particular review.

European guidelines<sup>139</sup> have noted that the procedure should be considered in carefully selected patients. We previously developed an approach in older adults that highlights physiologic as opposed to chronologic age<sup>140</sup>. A laparoscopic approach is favored as compared to an open approach. By applying the principles of a comprehensive geriatric assessment on patients evaluated for surgery, the hope is that those carefully selected individuals will have improved short- and long-term outcomes. Highlighting the importance of future life expectancy, presence of undiagnosed cognitive impairment, medical co-morbidity that could be impacted by the surgical procedure, and important social support mechanisms for the immediate post-operative care are of utmost importance. Previous history of post-operative delirium and impairments in vision and hearing are also important factors in successful recovery. Understanding such limitations could sway a decision to consider surgical intervention or not in a given patient. Being classified as 'geriatric' should

not preclude evaluation of surgery in those motivated older adults who fulfill many of the above noted elements in the geriatric evaluation.

## Conclusions

The epidemic of geriatric obesity will continue to impact the role of a primary care provider with time. The importance of lifespan prevention measures cannot be overstated to delay the onset of disability and impairments in health-related quality of life. Effective lifestyle modifications for weight loss can easily be implemented within a busy primary care setting to engage individuals. Community-based physical activity interventions are easy, cost-effective ways to delay disability and enhance physical function. Future studies should focus on disseminating and implementing practical ways to integrate established evidence-based practices into routine clinical care, without overburdening clinical staff. The use of emerging technologies may be helpful adjuncts. Evaluation of pharmacotherapy in this high-risk population remains a priority and including older, robust adults may be a reasonable first step in evaluating their safety and efficacy. We recommend that bariatric surgery be considered for older adults following a comprehensive geriatric assessment and interdisciplinary team-based approach is helping in evaluating and engaging these patients in this process.

## Acknowledgments

### Funding

Dr. Batsis' research reported in this publication was supported in part by the National Institute on Aging of the National Institutes of Health under Award Number K23AG051681. The content is solely the responsibility of the authors and does not necessarily represent the official views of the National Institutes of Health. This work was also supported by the Dartmouth Health Promotion and Disease Prevention Research Center (Cooperative Agreement Number U48DP005018) from the Centers for Disease Control and Prevention. The findings and conclusions in this journal article are those of the authors and do not necessarily represent the official position of the Centers for Disease Control and Prevention. Dr. Batsis has received honoraria from the Royal College of Physicians of Ireland for policy statement review and an honorarium from the Endocrine Society for an educational CME presentation at their Annual Conference.

## ABBREVIATIONS

<b>BMD</b>	bone mineral density
<b>BMI</b>	body mass index
<b>CI</b>	confidence intervals
<b>FNIH</b>	Foundation for the National Institute on Health
<b>HR</b>	hazard ratio
<b>NHANES</b>	National Health and Nutrition and Examination Survey

## References

1. Government US. Census Bureau Statistics. 2012 Accessed January 20th, 2013, 2013.
2. Mather, M., Jacobsen, LA., Pollard, KM. Aging in the United States. Population Reference Bureau; 2015. Available at: <http://www.prb.org/pdf16/aging-us-population-bulletin.pdf>

3. Lubitz J, Cai L, Kramarow E, Lentzner H. Health, life expectancy, and health care spending among the elderly. *N Engl J Med*. Sep 11; 2003 349(11):1048–1055. [PubMed: 12968089]
4. Katz S, Branch LG, Branson MH, Papsidero JA, Beck JC, Greer DS. Active life expectancy. *N Engl J Med*. Nov 17; 1983 309(20):1218–1224. [PubMed: 6633571]
5. Arias E. National Vital Statistics Report. Sep 22.2015 64(11)
6. Dunlop DD, Hughes SL, Manheim LM. Disability in activities of daily living: patterns of change and a hierarchy of disability. *Am J Public Health*. Mar; 1997 87(3):378–383. [PubMed: 9096537]
7. Katz S, Ford AB, Moskowitz RW, Jackson BA, Jaffe MW. Studies of Illness in the Aged. The Index of Adl: A Standardized Measure of Biological and Psychosocial Function. *JAMA*. Sep 21.1963 185:914–919. [PubMed: 14044222]
8. Bish CL, Michels Blanck H, Maynard LM, Serdula MK, Thompson NJ, Kettel Khan L. Health-related quality of life and weight loss among overweight and obese U.S. adults, 2001 to 2002. *Obesity (Silver Spring)*. Nov; 2006 14(11):2042–2053. [PubMed: 17135622]
9. Cai Q, Salmon JW, Rodgers ME. Factors associated with long-stay nursing home admissions among the U.S. elderly population: comparison of logistic regression and the Cox proportional hazards model with policy implications for social work. *Soc Work Health Care*. 2009; 48(2):154–168. [PubMed: 19197772]
10. Chambers BA, Guo SS, Siervogel R, Hall G, Chumlea WC. Cumulative effects of cardiovascular disease risk factors on quality of life. *J Nutr Health Aging*. May; 2002 6(3):179–184. [PubMed: 11887243]
11. Cigolle CT, Langa KM, Kabeto MU, Tian Z, Blaum CS. Geriatric conditions and disability: the Health and Retirement Study. *Ann Intern Med*. Aug 07; 2007 147(3):156–164. [PubMed: 17679703]
12. Flegal KM, Kruszon-Moran D, Carroll MD, Fryar CD, Ogden CL. Trends in Obesity Among Adults in the United States, 2005 to 2014. *JAMA*. Jun 07; 2016 315(21):2284–2291. [PubMed: 27272580]
13. Baker C. Obesity Statistics: A briefing Paper. House of Commons Library. Jan 20.2017 3336
14. Twells LK, Gregory DM, Reddigan J, Midodzi WK. Current and predicted prevalence of obesity in Canada: a trend analysis. *CMAJ Open*. Jan; 2014 2(1):E18–26.
15. Baumgartner RN. Body composition in healthy aging. *Ann N Y Acad Sci*. May.2000 904:437–448. [PubMed: 10865787]
16. Sayer AA, Syddall H, Martin H, Patel H, Baylis D, Cooper C. The developmental origins of sarcopenia. *J Nutr Health Aging*. Aug-Sep;2008 12(7):427–432. [PubMed: 18615224]
17. Fielding RA, Vellas B, Evans WJ, et al. Sarcopenia: an undiagnosed condition in older adults. Current consensus definition: prevalence, etiology, and consequences. International working group on sarcopenia. *J Am Med Dir Assoc*. May; 2011 12(4):249–256. [PubMed: 21527165]
18. Dutta C. Significance of sarcopenia in the elderly. *J Nutr*. May; 1997 127(5 Suppl):992S–993S. [PubMed: 9164281]
19. Bales CW, Ritchie CS. Sarcopenia, weight loss, and nutritional frailty in the elderly. *Annu Rev Nutr*. 2002; 22:309–323. [PubMed: 12055348]
20. Studenski SA, Peters KW, Alley DE, et al. The FNIH sarcopenia project: rationale, study description, conference recommendations, and final estimates. *J Gerontol A Biol Sci Med Sci*. May; 2014 69(5):547–558. [PubMed: 24737557]
21. Sorkin JD, Muller DC, Andres R. Longitudinal change in height of men and women: implications for interpretation of the body mass index: the Baltimore Longitudinal Study of Aging. *Am J Epidemiol*. Nov 01; 1999 150(9):969–977. [PubMed: 10547143]
22. Romero-Corral A, Somers VK, Sierra-Johnson J, et al. Accuracy of body mass index in diagnosing obesity in the adult general population. *Int J Obes (Lond)*. Jun; 2008 32(6):959–966. [PubMed: 18283284]
23. Okorodudu DO, Jumean MF, Montori VM, et al. Diagnostic performance of body mass index to identify obesity as defined by body adiposity: a systematic review and meta-analysis. *Int J Obes (Lond)*. May; 2010 34(5):791–799. [PubMed: 20125098]

24. Batsis JA, Mackenzie TA, Bartels SJ, Sahakyan KR, Somers VK, Lopez-Jimenez F. Diagnostic accuracy of body mass index to identify obesity in older adults: NHANES 1999–2004. *Int J Obes (Lond)*. May; 2016 40(5):761–767. [PubMed: 26620887]
25. Batsis JA, Zbehlik AJ, Scherer EA, Barre LK, Bartels SJ. Normal Weight with Central Obesity, Physical Activity, and Functional Decline: Data from the Osteoarthritis Initiative. *J Am Geriatr Soc*. Aug; 2015 63(8):1552–1560. [PubMed: 26173812]
26. Romero-Corral A, Somers VK, Sierra-Johnson J, et al. Normal weight obesity: a risk factor for cardiometabolic dysregulation and cardiovascular mortality. *Eur Heart J*. Mar; 2010 31(6):737–746. [PubMed: 19933515]
27. Batsis JA, Sahakyan KR, Rodriguez-Escudero JP, Bartels SJ, Somers VK, Lopez-Jimenez F. Normal weight obesity and mortality in United States subjects  $\geq 60$  years of age (from the Third National Health and Nutrition Examination Survey). *Am J Cardiol*. Nov 15; 2013 112(10):1592–1598. [PubMed: 23993123]
28. Batsis JA, Sahakyan KR, Rodriguez-Escudero JP, Bartels SJ, Lopez-Jimenez F. Normal weight obesity and functional outcomes in older adults. *Eur J Intern Med*. Jul; 2014 25(6):517–522. [PubMed: 24909976]
29. Sahakyan KR, Somers VK, Rodriguez-Escudero JP, et al. Normal-Weight Central Obesity: Implications for Total and Cardiovascular Mortality. *Ann Intern Med*. Dec 01; 2015 163(11):827–835. [PubMed: 26551006]
30. Schlanger LE, Bailey JL, Sands JM. Electrolytes in the aging. *Adv Chronic Kidney Dis*. Jul; 2010 17(4):308–319. [PubMed: 20610358]
31. Maradit Kremers H, Larson DR, Crowson CS, et al. Prevalence of Total Hip and Knee Replacement in the United States. *J Bone Joint Surg Am*. Sep 02; 2015 97(17):1386–1397. [PubMed: 26333733]
32. Silverman BG, Gross TP, Kaczmarek RG, Hamilton P, Hamburger S. The epidemiology of pacemaker implantation in the United States. *Public Health Rep*. Jan-Feb; 1995 110(1):42–46. [PubMed: 7838942]
33. Flegal KM, Kit BK, Orpana H, Graubard BI. Association of all-cause mortality with overweight and obesity using standard body mass index categories: a systematic review and meta-analysis. *JAMA*. Jan 02; 2013 309(1):71–82. [PubMed: 23280227]
34. Flegal KM, Graubard BI, Williamson DF, Gail MH. Excess deaths associated with underweight, overweight, and obesity. *JAMA*. Apr 20; 2005 293(15):1861–1867. [PubMed: 15840860]
35. Houston DK, Ding J, Nicklas BJ, et al. Overweight and obesity over the adult life course and incident mobility limitation in older adults: the health, aging and body composition study. *Am J Epidemiol*. Apr 15; 2009 169(8):927–936. [PubMed: 19270048]
36. Strandberg TE, Sirola J, Pitkala KH, Tilvis RS, Strandberg AY, Stenholm S. Association of midlife obesity and cardiovascular risk with old age frailty: a 26-year follow-up of initially healthy men. *Int J Obes (Lond)*. Sep; 2012 36(9):1153–1157. [PubMed: 22614054]
37. Elkins JS, Whitmer RA, Sidney S, Sorel M, Yaffe K, Johnston SC. Midlife obesity and long-term risk of nursing home admission. *Obesity (Silver Spring)*. Aug; 2006 14(8):1472–1478. [PubMed: 16988091]
38. Winter JE, MacInnis RJ, Wattanapenpaiboon N, Nowson CA. BMI and all-cause mortality in older adults: a meta-analysis. *Am J Clin Nutr*. Apr; 2014 99(4):875–890. [PubMed: 24452240]
39. Veronese N, Cereda E, Solmi M, et al. Inverse relationship between body mass index and mortality in older nursing home residents: a meta-analysis of 19,538 elderly subjects. *Obes Rev*. Nov; 2015 16(11):1001–1015. [PubMed: 26252230]
40. Lee CD, Blair SN, Jackson AS. Cardiorespiratory fitness, body composition, and all-cause and cardiovascular disease mortality in men. *Am J Clin Nutr*. Mar; 1999 69(3):373–380. [PubMed: 10075319]
41. McAuley PA, Artero EG, Sui X, et al. The obesity paradox, cardiorespiratory fitness, and coronary heart disease. *Mayo Clin Proc*. May; 2012 87(5):443–451. [PubMed: 22503065]
42. Schaap LA, Koster A, Visser M. Adiposity, muscle mass, and muscle strength in relation to functional decline in older persons. *Epidemiol Rev*. Dec 4. 2013 35:51–65. [PubMed: 23221972]

43. Zizza CA, Herring A, Stevens J, Popkin BM. Obesity affects nursing-care facility admission among whites but not blacks. *Obes Res.* Aug; 2002 10(8):816–823. [PubMed: 12181391]
44. Chang VW, Alley DE, Dowd JB. Trends in the Relationship of Obesity and Disability, 1988–2012. *Am J Epidemiol.* May 09.2017
45. Visser M, Langlois J, Guralnik JM, et al. High body fatness, but not low fat-free mass, predicts disability in older men and women: the Cardiovascular Health Study. *Am J Clin Nutr.* Sep; 1998 68(3):584–590. [PubMed: 9734734]
46. Visser M, Harris TB, Langlois J, et al. Body fat and skeletal muscle mass in relation to physical disability in very old men and women of the Framingham Heart Study. *J Gerontol A Biol Sci Med Sci.* May; 1998 53(3):M214–221. [PubMed: 9597054]
47. Masud T, Morris RO. Epidemiology of falls. *Age Ageing.* Nov; 2001 30(Suppl 4):3–7.
48. Tinetti ME. Clinical practice. Preventing falls in elderly persons. *N Engl J Med.* Jan 02; 2003 348(1):42–49. [PubMed: 12510042]
49. American Geriatrics Society Workgroup on Vitamin DSfOA. Recommendations abstracted from the American Geriatrics Society Consensus Statement on vitamin D for Prevention of Falls and Their Consequences. *J Am Geriatr Soc.* Jan; 2014 62(1):147–152. [PubMed: 24350602]
50. Himes CL, Reynolds SL. Effect of obesity on falls, injury, and disability. *J Am Geriatr Soc.* Jan; 2012 60(1):124–129. [PubMed: 22150343]
51. Ylitalo KR, Karvonen-Gutierrez CA. Body mass index, falls, and injurious falls among U.S. adults: Findings from the 2014 Behavioral Risk Factor Surveillance System. *Prev Med.* Oct.2016 91:217–223. [PubMed: 27575319]
52. Tang X, Liu G, Kang J, et al. Obesity and risk of hip fracture in adults: a meta-analysis of prospective cohort studies. *PLoS One.* 2013; 8(4):e55077. [PubMed: 23593112]
53. Delmonico MJ, Harris TB, Visser M, et al. Longitudinal study of muscle strength, quality, and adipose tissue infiltration. *Am J Clin Nutr.* Dec; 2009 90(6):1579–1585. [PubMed: 19864405]
54. Visser M, Kritchevsky SB, Goodpaster BH, et al. Leg muscle mass and composition in relation to lower extremity performance in men and women aged 70 to 79: the health, aging and body composition study. *J Am Geriatr Soc.* May; 2002 50(5):897–904. [PubMed: 12028178]
55. Batsis JA, Barre LK, Mackenzie TA, Pratt SI, Lopez-Jimenez F, Bartels SJ. Variation in the prevalence of sarcopenia and sarcopenic obesity in older adults associated with different research definitions: dual-energy X-ray absorptiometry data from the National Health and Nutrition Examination Survey 1999–2004. *J Am Geriatr Soc.* Jun; 2013 61(6):974–980. [PubMed: 23647372]
56. Clark BC, Manini TM. Functional consequences of sarcopenia and dynapenia in the elderly. *Curr Opin Clin Nutr Metab Care.* May; 2010 13(3):271–276. [PubMed: 20154609]
57. Menant JC, Weber F, Lo J, et al. Strength measures are better than muscle mass measures in predicting health-related outcomes in older people: time to abandon the term sarcopenia? *Osteoporos Int.* Jan; 2017 28(1):59–70. [PubMed: 27394415]
58. Villareal DT, Chode S, Parimi N, et al. Weight loss, exercise, or both and physical function in obese older adults. *N Engl J Med.* Mar 31; 2011 364(13):1218–1229. [PubMed: 21449785]
59. Goodpaster BH, Park SW, Harris TB, et al. The loss of skeletal muscle strength, mass, and quality in older adults: the health, aging and body composition study. *J Gerontol A Biol Sci Med Sci.* Oct; 2006 61(10):1059–1064. [PubMed: 17077199]
60. Baumgartner RN, Wayne SJ, Waters DL, Janssen I, Gallagher D, Morley JE. Sarcopenic obesity predicts instrumental activities of daily living disability in the elderly. *Obes Res.* Dec; 2004 12(12):1995–2004. [PubMed: 15687401]
61. Baumgartner RN, Koehler KM, Gallagher D, et al. Epidemiology of sarcopenia among the elderly in New Mexico. *Am J Epidemiol.* Apr; 1998 147(8):755–763. [PubMed: 9554417]
62. Stenholm S, Alley D, Bandinelli S, et al. The effect of obesity combined with low muscle strength on decline in mobility in older persons: results from the InCHIANTI study. *Int J Obes (Lond).* Jun; 2009 33(6):635–644. [PubMed: 19381155]
63. Batsis JA, Mackenzie TA, Lopez-Jimenez F, Bartels SJ. Sarcopenia, sarcopenic obesity, and functional impairments in older adults: National Health and Nutrition Examination Surveys 1999–2004. *Nutr Res.* Dec; 2015 35(12):1031–1039. [PubMed: 26472145]



64. Batsis JA, Mackenzie TA, Emeny RT, Lopez-Jimenez F, Bartels SJ. Low Lean Mass With and Without Obesity, and Mortality: Results From the 1999–2004 National Health and Nutrition Examination Survey. *J Gerontol A Biol Sci Med Sci*. Feb 16.2017
65. Batsis JA, Mackenzie TA, Barre LK, Lopez-Jimenez F, Bartels SJ. Sarcopenia, sarcopenic obesity and mortality in older adults: results from the National Health and Nutrition Examination Survey III. *Eur J Clin Nutr*. Sep; 2014 68(9):1001–1007. [PubMed: 24961545]
66. Hardy R, Kuh D. Commentary: BMI and mortality in the elderly—a life course perspective. *Int J Epidemiol*. Feb; 2006 35(1):179–180. [PubMed: 16394115]
67. Richman EL, Stampfer MJ. Weight loss and mortality in the elderly: separating cause and effect. *J Intern Med*. Aug; 2010 268(2):103–105. [PubMed: 20642755]
68. Villareal DT, Apovian CM, Kushner RF, Klein S, American Society for N, Naaso TOS. Obesity in older adults: technical review and position statement of the American Society for Nutrition and NAASO, The Obesity Society. *Am J Clin Nutr*. Nov; 2005 82(5):923–934. [PubMed: 16280421]
69. Batsis JA, Gill LE, Masutani RK, et al. Weight Loss Interventions in Older Adults with Obesity: A Systematic Review of Randomized Controlled Trials Since 2005. *J Am Geriatr Soc*. Feb; 2017 65(2):257–268. [PubMed: 27641543]
70. Jacob ME, Yee LM, Diehr PH, et al. Can a Healthy Lifestyle Compress the Disabled Period in Older Adults? *J Am Geriatr Soc*. Oct; 2016 64(10):1952–1961. [PubMed: 27603679]
71. Villareal DT, Aguirre L, Gurney AB, et al. Aerobic or Resistance Exercise, or Both in Dieting Obese Older Adults. *N Engl J Med*. 2017; 376:1943–1955. [PubMed: 28514618]
72. Kritchevsky SB, Beavers KM, Miller ME, et al. Intentional weight loss and all-cause mortality: a meta-analysis of randomized clinical trials. *PLoS One*. 2015; 10(3):e0121993. [PubMed: 25794148]
73. Gill LE, Bartels SJ, Batsis JA. Weight Management in Older Adults. *Curr Obes Rep*. Sep; 2015 4(3):379–388. [PubMed: 26627496]
74. Heymsfield SB, Gonzalez MC, Shen W, Redman L, Thomas D. Weight loss composition is one-fourth fat-free mass: a critical review and critique of this widely cited rule. *Obes Rev*. Apr; 2014 15(4):310–321. [PubMed: 24447775]
75. Soltani S, Hunter GR, Kazemi A, Shab-Bidar S. The effects of weight loss approaches on bone mineral density in adults: a systematic review and meta-analysis of randomized controlled trials. *Osteoporos Int*. Sep; 2016 27(9):2655–2671. [PubMed: 27154437]
76. Zibellini J, Seimon RV, Lee CM, Gibson AA, Hsu MS, Sainsbury A. Effect of diet-induced weight loss on muscle strength in adults with overweight or obesity – a systematic review and meta-analysis of clinical trials. *Obes Rev*. Aug; 2016 17(8):647–663. [PubMed: 27126087]
77. Zibellini J, Seimon RV, Lee CM, et al. Does Diet-Induced Weight Loss Lead to Bone Loss in Overweight or Obese Adults? A Systematic Review and Meta-Analysis of Clinical Trials. *J Bone Miner Res*. Dec; 2015 30(12):2168–2178. [PubMed: 26012544]
78. Shah K, Armamento-Villareal R, Parimi N, et al. Exercise training in obese older adults prevents increase in bone turnover and attenuates decrease in hip bone mineral density induced by weight loss despite decline in bone-active hormones. *J Bone Miner Res*. Dec; 2011 26(12):2851–2859. [PubMed: 21786319]
79. Tinetti ME, Doucette J, Claus E, Marottoli R. Risk factors for serious injury during falls by older persons in the community. *J Am Geriatr Soc*. Nov; 1995 43(11):1214–1221. [PubMed: 7594154]
80. Batsis JA, Huddleston JM, Melton LJ, et al. Body mass index and risk of adverse cardiac events in elderly patients with hip fracture: a population-based study. *J Am Geriatr Soc*. Mar; 2009 57(3):419–426. [PubMed: 19175436]
81. Wadden TA, Butryn ML, Hong PS, Tsai AG. Behavioral treatment of obesity in patients encountered in primary care settings: a systematic review. *JAMA*. Nov 05; 2014 312(17):1779–1791. [PubMed: 25369490]
82. Tsai AG, Wadden TA. Treatment of obesity in primary care practice in the United States: a systematic review. *J Gen Intern Med*. Sep; 2009 24(9):1073–1079. [PubMed: 19562419]
83. Aveyard P, Lewis A, Tearne S, et al. Screening and brief intervention for obesity in primary care: a parallel, two-arm, randomised trial. *Lancet*. Nov 19; 2016 388(10059):2492–2500. [PubMed: 27789061]

84. Decision Memo for Intensive Behavioral Therapy for Obesity (CAG-00423N). Centers for Medicare & Medicaid Services; 2011. <http://www.cms.gov/medicare-coverage-database/details/nca-decision-memo.aspx?&NcaName=Intensive&percent;20Behavioral%20Therapy%20for%20Obesity&bc=ACAAAAAIAAA&NCAId=253>. Accessed March 22nd, 2013
85. Batsis JA, Huyck KL, Bartels SJ. Challenges with the Medicare obesity benefit: practical concerns & proposed solutions. *J Gen Intern Med.* Jan; 2015 30(1):118–122. [PubMed: 25227742]
86. Batsis JA, Pletcher SN, Stahl JE. Telemedicine and primary care obesity management in rural areas – innovative approach for older adults? *BMC Geriatr.* Jan 05.2017 17(1):6. [PubMed: 28056832]
87. Batsis JA, Bynum JP. Uptake of the centers for medicare and medicaid obesity benefit: 2012–2013. *Obesity (Silver Spring).* Sep; 2016 24(9):1983–1988. [PubMed: 27465909]
88. Waters DL, Baumgartner RN, Garry PJ, Vellas B. Advantages of dietary, exercise-related, and therapeutic interventions to prevent and treat sarcopenia in adult patients: an update. *Clin Interv Aging.* Sep 07.2010 5:259–270. [PubMed: 20852673]
89. Jensen MD, Ryan DH, Apovian CM, et al. 2013 AHA/ACC/TOS Guideline for the Management of Overweight and Obesity in Adults. *Circulation.* Jun 24; 2014 129(25 suppl 2):S102–S138. [PubMed: 24222017]
90. De Luca, CR., Leventer, RJ. Developmental trajectories of executive functions across the lifespan. Washington, DC: Taylor & Francis; 2008.
91. Ende J, Kazis L, Ash A, Moskowitz MA. Measuring patients' desire for autonomy: decision making and information-seeking preferences among medical patients. *J Gen Intern Med.* Jan-Feb; 1989 4(1):23–30. [PubMed: 2644407]
92. Orbell S, Sheeran P. 'Inclined abstainers': a problem for predicting health-related behaviour. *Br J Soc Psychol.* Jun; 1998 37(Pt 2):151–165. [PubMed: 9639861]
93. Allan JL, Sniehotta FF, Johnston M. The best laid plans: planning skill determines the effectiveness of action plans and implementation intentions. *Ann Behav Med.* Aug; 2013 46(1):114–120. [PubMed: 23456214]
94. Allan JL, Johnston M, Campbell N. Missed by an inch or a mile? Predicting the size of intention-behaviour gap from measures of executive control. *Psychol Health.* Jun; 2011 26(6):635–650. [PubMed: 21360414]
95. Nigg CR, Long CR. A systematic review of single health behavior change interventions vs. multiple health behavior change interventions among older adults. *Transl Behav Med.* Jun; 2012 2(2):163–179. [PubMed: 24073109]
96. French DP, Olander EK, Chisholm A, Mc Sharry J. Which behaviour change techniques are most effective at increasing older adults' self-efficacy and physical activity behaviour? A systematic review. *Ann Behav Med.* Oct; 2014 48(2):225–234. [PubMed: 24648017]
97. Purath J, Keck A, Fitzgerald CE. Motivational interviewing for older adults in primary care: a systematic review. *Geriatr Nurs.* May-Jun;2014 35(3):219–224. [PubMed: 24656051]
98. Ziegelmann JP, Knoll N. Future Directions in the Study of Health Behavior among Older Adults. *Gerontology.* 2015; 61(5):469–476. [PubMed: 25660128]
99. Lum TY, Lightfoot E. The effects of volunteering on the physical and mental health of older people. *Res Aging.* 2005; 27:31–55.
100. Guiding principles for the care of older adults with multimorbidity: an approach for c. Guiding principles for the care of older adults with multimorbidity: an approach for clinicians: American Geriatrics Society Expert Panel on the Care of Older Adults with Multimorbidity. *J Am Geriatr Soc.* Oct; 2012 60(10):E1–E25. [PubMed: 22994865]
101. Freund AM. Age-differential motivational consequences of optimization versus compensation focus in younger and older adults. *Psychol Aging.* Jun; 2006 21(2):240–252. [PubMed: 16768572]
102. Levy BR, Slade MD, Kasl SV. Longitudinal benefit of positive self-perceptions of aging on functional health. *J Gerontol B Psychol Sci Soc Sci.* Sep; 2002 57(5):P409–417. [PubMed: 12198099]
103. Lockenhoff CE, Carstensen LL. Socioemotional selectivity theory, aging, and health: the increasingly delicate balance between regulating emotions and making tough choices. *J Pers.* Dec; 2004 72(6):1395–1424. [PubMed: 15509287]

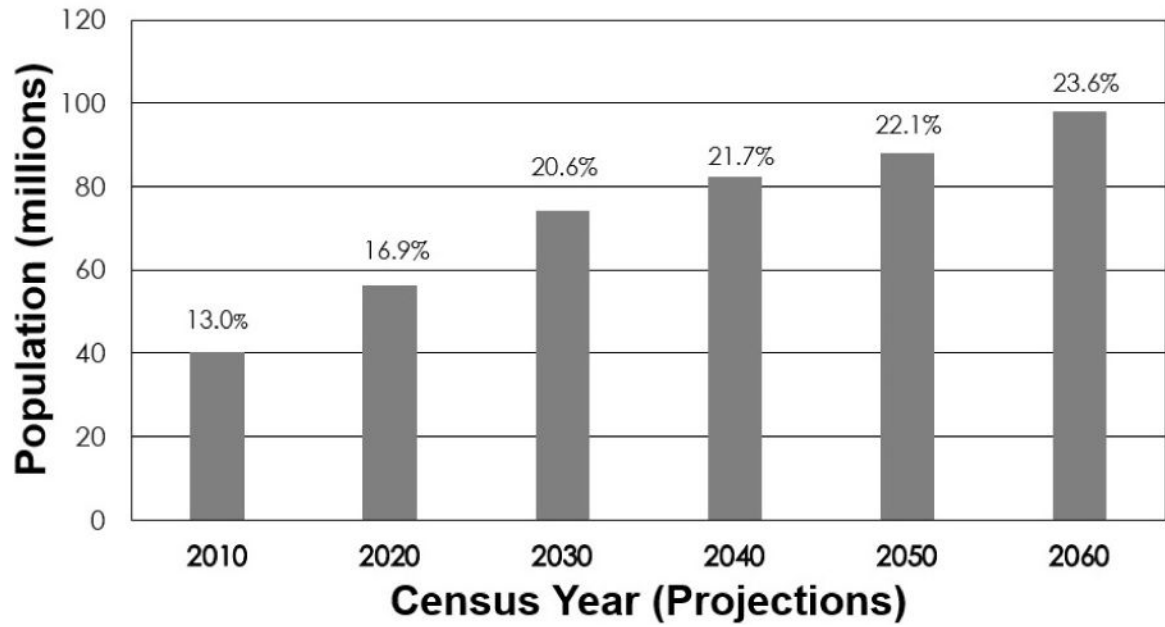
104. Newsom JT, Huguet N, McCarthy MJ, et al. Health behavior change following chronic illness in middle and later life. *J Gerontol B Psychol Sci Soc Sci*. May; 2012 67(3):279–288. [PubMed: 21983040]
105. Keller H, Laporte M, Payette H, et al. Prevalence and predictors of weight change post discharge from hospital: a study of the Canadian Malnutrition Task Force. *Eur J Clin Nutr*. Feb 22.2017
106. Fulton MM, Allen ER. Polypharmacy in the elderly: a literature review. *J Am Acad Nurse Pract*. Apr; 2005 17(4):123–132. [PubMed: 15819637]
107. American Geriatrics Society. 2015 Updated Beers Criteria for Potentially Inappropriate Medication Use in Older Adults. *J Am Geriatr Soc*. Nov; 2015 63(11):2227–2246. [PubMed: 26446832]
108. Kirkman MS, Briscoe VJ, Clark N, et al. Diabetes in older adults: a consensus report. *J Am Geriatr Soc*. Dec; 2012 60(12):2342–2356. [PubMed: 23106132]
109. Soysal P, Veronese N, Thompson T, et al. Relationship between depression and frailty in older adults: A systematic review and meta-analysis. *Ageing Res Rev*. Mar 31.2017 36:78–87. [PubMed: 28366616]
110. Sibille KT, McBeth J, Smith D, Wilkie R. Allostatic load and pain severity in older adults: Results from the English Longitudinal Study of Ageing. *Exp Gerontol*. Feb.2017 88:51–58. [PubMed: 27988258]
111. Mendelson G, Ness J, Aronow WS. Drug treatment of hypertension in older persons in an academic hospital-based geriatrics practice. *J Am Geriatr Soc*. May; 1999 47(5):597–599. [PubMed: 10323654]
112. Castillo DC, Ramsey NLM, Yu SSK, Ricks M, Courville AB, Sumner AE. Inconsistent Access to Food and Cardiometabolic Disease: The Effect of Food Insecurity. *Curr Cardiovasc Risk Rep*. 2012; 6(3):245–250. [PubMed: 22629473]
113. Lee JS, Frongillo EA. Nutritional and Health Consequences Are Associated with Food Insecurity among U.S. Elderly Persons. *J Nutr*. 2001; 131:1503–1509. [PubMed: 11340107]
114. Appel LJ, Moore TJ, Obarzanek E, et al. A clinical trial of the effects of dietary patterns on blood pressure. DASH Collaborative Research Group. *N Engl J Med*. Apr 17; 1997 336(16):1117–1124. [PubMed: 9099655]
115. Esposito K, Kastorini CM, Panagiotakos DB, Giugliano D. Mediterranean diet and weight loss: meta-analysis of randomized controlled trials. *Metab Syndr Relat Disord*. Feb; 2011 9(1):1–12. [PubMed: 20973675]
116. Bauer J, Biolo G, Cederholm T, et al. Evidence-based recommendations for optimal dietary protein intake in older people: a position paper from the PROT-AGE Study Group. *J Am Med Dir Assoc*. Aug; 2013 14(8):542–559. [PubMed: 23867520]
117. Colonetti T, Grande AJ, Milton K, et al. Effects of whey protein supplement in the elderly submitted to resistance training: systematic review and meta-analysis. *Int J Food Sci Nutr*. May; 2017 68(3):257–264. [PubMed: 27653283]
118. Porter Starr KN, Pieper CF, Orenduff MC, et al. Improved Function With Enhanced Protein Intake per Meal: A Pilot Study of Weight Reduction in Frail, Obese Older Adults. *J Gerontol A Biol Sci Med Sci*. Oct; 2016 71(10):1369–1375. [PubMed: 26786203]
119. Carroll JK, Antognoli E, Flocke SA. Evaluation of physical activity counseling in primary care using direct observation of the 5As. *Ann Fam Med*. Sep-Oct;2011 9(5):416–422. [PubMed: 21911760]
120. Garber CE, Blissmer B, Deschenes MR, et al. American College of Sports Medicine position stand. Quantity and quality of exercise for developing and maintaining cardiorespiratory, musculoskeletal, and neuromotor fitness in apparently healthy adults: guidance for prescribing exercise. *Med Sci Sports Exerc*. Jul; 2011 43(7):1334–1359. [PubMed: 21694556]
121. Pahor M, Guralnik JM, Ambrosius WT, et al. Effect of structured physical activity on prevention of major mobility disability in older adults: the LIFE study randomized clinical trial. *JAMA*. Jun 18; 2014 311(23):2387–2396. [PubMed: 24866862]
122. Screening for osteoporosis: U.S. preventive services task force recommendation statement. *Ann Intern Med*. Mar 01; 2011 154(5):356–364. [PubMed: 21242341]

123. Garvey WT, Mechanick JI, Brett EM, et al. American Association of Clinical Endocrinologists and American College of Endocrinology Comprehensive Clinical Practice Guidelines for Medical Care of Patients with Obesity. *Endocr Pract.* Jul; 2016 22(Suppl 3):1–203.
124. Davies MJ, Bergenstal R, Bode B, et al. Efficacy of Liraglutide for Weight Loss Among Patients With Type 2 Diabetes: The SCALE Diabetes Randomized Clinical Trial. *JAMA.* Aug 18; 2015 314(7):687–699. [PubMed: 26284720]
125. Gadde KM, Allison DB, Ryan DH, et al. Effects of low-dose, controlled-release, phentermine plus topiramate combination on weight and associated comorbidities in overweight and obese adults (CONQUER): a randomised, placebo-controlled, phase 3 trial. *Lancet.* Apr 16; 2011 377(9774):1341–1352. [PubMed: 21481449]
126. Perna S, Guido D, Bologna C, et al. Liraglutide and obesity in elderly: efficacy in fat loss and safety in order to prevent sarcopenia. A perspective case series study. *Aging Clin Exp Res.* Dec; 2016 28(6):1251–1257. [PubMed: 26749118]
127. Pi-Sunyer X, Astrup A, Fujioka K, et al. A Randomized, Controlled Trial of 3.0 mg of Liraglutide in Weight Management. *N Engl J Med.* Jul 02; 2015 373(1):11–22. [PubMed: 26132939]
128. Wadden TA, Hollander P, Klein S, et al. Weight maintenance and additional weight loss with liraglutide after low-calorie-diet-induced weight loss: the SCALE Maintenance randomized study. *Int J Obes (Lond).* Nov; 2013 37(11):1443–1451. [PubMed: 23812094]
129. Gastrointestinal surgery for severe obesity. *Am J Clin Nutr*; Proceedings of a National Institutes of Health Consensus Development Conference; March 25–27, 1991; Bethesda, MD. Feb. 1992 p. 487s-619s.
130. Meron Eldar SHH, Kroh M, Chand B, Rogula T, Schauer P, Brethauer S. Bariatric surgery in the elderly – Comparing the 3 major bariatric procedures. *Obes Surg.* 2011; 21(8):1134–1135.
131. Batsis JA, Miranda WR, Prasad C, et al. Effect of bariatric surgery on cardiometabolic risk in elderly patients: A population-based study. *Geriatr Gerontol Int.* May; 2016 16(5):618–624. [PubMed: 26017642]
132. Flum DR, Kwon S, MacLeod K, et al. The use, safety and cost of bariatric surgery before and after Medicare’s national coverage decision. *Ann Surg.* Dec; 2011 254(6):860–865. [PubMed: 21975317]
133. O’Keefe KL, Kemmeter PR, Kemmeter KD. Bariatric surgery outcomes in patients aged 65 years and older at an American Society for Metabolic and Bariatric Surgery Center of Excellence. *Obes Surg.* Sep; 2010 20(9):1199–1205. [PubMed: 20532834]
134. Quebbemann B, Engstrom D, Siegfried T, Garner K, Dallal R. Bariatric surgery in patients older than 65 years is safe and effective. *Surg Obes Relat Dis.* Jul-Aug;2005 1(4):389–392. discussion 392–383. [PubMed: 16925254]
135. Ramirez A, Roy M, Hidalgo JE, Szomstein S, Rosenthal RJ. Outcomes of bariatric surgery in patients >70 years old. *Surg Obes Relat Dis.* Jul-Aug;2012 8(4):458–462. [PubMed: 22551574]
136. Ibrahim AM, Ghaferi AA, Thumma JR, Dimick JB. Variation in Outcomes at Bariatric Surgery Centers of Excellence. *JAMA Surg.* Apr 26.2017
137. Livingston EH. Bariatric surgery outcomes at designated centers of excellence vs nondesignated programs. *Arch Surg.* Apr; 2009 144(4):319–325. discussion 325. [PubMed: 19380644]
138. Giordano S, Victorzon M. Bariatric surgery in elderly patients: a systematic review. *Clin Interv Aging.* 2015; 10:1627–1635. [PubMed: 26508845]
139. Matus-Vliegen EM, Obesity Management Task Force of the European Association for the Study of O. Prevalence, pathophysiology, health consequences and treatment options of obesity in the elderly: a guideline. *Obes Facts.* 2012; 5(3):460–483. [PubMed: 22797374]
140. Batsis JA, Dolkart KM. Evaluation of older Adults with obesity for bariatric surgery: Geriatricians’ perspective. *Journal of Clinical Gerontology and Geriatrics.* 2015; 6(2):45–53.

**Key Points**

- Older adults with obesity will be an emerging demographic that primary care practitioners will need to develop skills in managing.
- Intentional weight loss in this population can be successful and safe.
- Appropriate understanding of the dangers of weight loss on muscle and bone are required.
- Pharmacotherapies that are FDA approved for adults have not been extensively studied in older adult populations.
- Bariatric surgery can be considered in selective candidates.

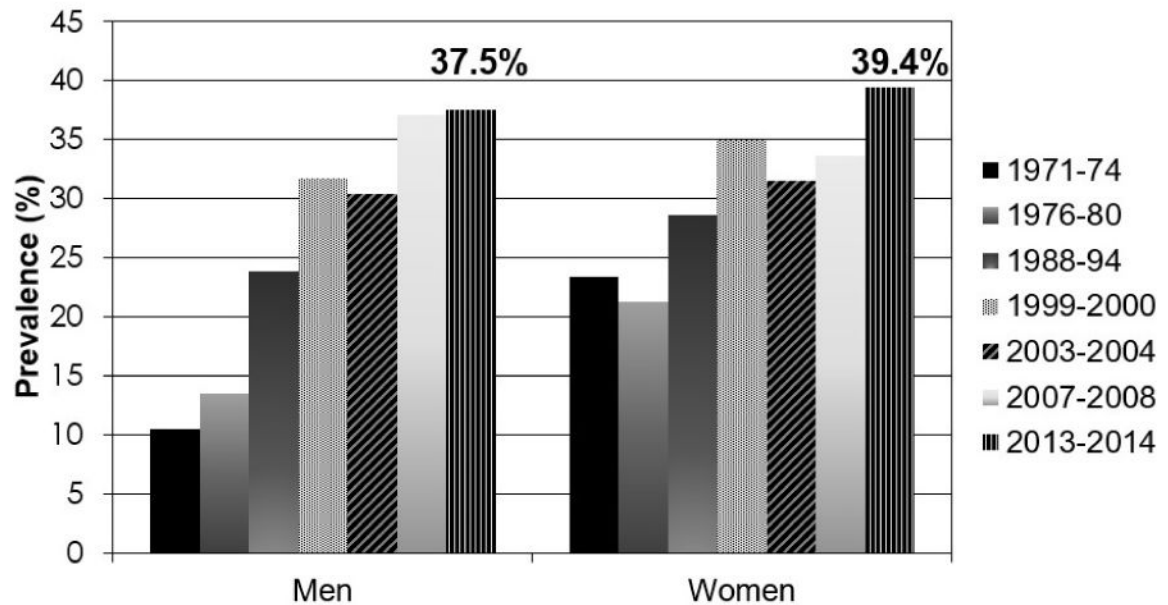
## Projected Elderly Population ( $\geq 65$ years) United States, 2010-2050



**Figure 1. Projected Elderly Population Aged 65 years and older (represented as percentages) in the United States based on the 2010 Census and Future Projections**

Data from United States Census Bureau. 2010 Census Data. Available at: <https://www.census.gov/2010census/data/>. Accessed Jan 20 2013.

## Prevalence of Obesity in Ages 60+ National Health & Nutrition Examination Surveys 1971-2014



**Figure 2. Prevalence of Obesity in ages 60 years and older using National Health and Nutrition Examination Surveys over the past 4 decades based on body mass index. Current estimates indicate that obesity is present in men in 37.5% and 39.4% in females in the population**  
*Data from Flegal KM, Kruszon-Moran D, Carroll MD, et al. Trends in Obesity Among Adults in the United States, 2005 to 2014. JAMA 2016;315(21):2284–2291.*

**Table 1**

Commonly Prescribed Medications in Older Adults Predisposing to Weight Gain

Disease/Class of Med	Examples
Diabetes	Insulin, TDZ *, sulfonylureas *
Depression	Tricyclics *, SSRIs *
Antipsychotics	1 <sup>st</sup> + 2 <sup>nd</sup> generation *
Neuropathic	gabapentin
Antihistamines	Diphenhydramine *
Hypertension	B-blockers

\* Medications that are listed in the 2015 Beers Criteria<sup>107</sup>

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript



**Table 2**

Relative and Absolute Contraindications to Weight-Loss Medications in Older Adults

<b>Generic Name</b>	<b>Absolute Contraindications</b>	<b>Side-Effects (&gt;10%) &amp; Major Risks in Older Adults</b>
<b>Locaserin</b>	—	Renal insufficiency, Tramadol, Heart failure, Serotonin excess, hypoglycemia
<b>Phentermine/Topiramate</b>	Glaucoma, MAOI hyperthyroidism	Constipation headache, xerostomia
<b>Phentermine</b>	Glaucoma, Heart failure, <b>CAD</b> , Hyperthyroidism, <b>Arrhythmias</b>	Renal insufficiency, Reduced exercise tolerance
<b>Orlistat</b>	Malabsorption, cholestasis	Fecal urgency, Flatulence, Steatorrhea
<b>Bupropion/naltrexone</b>	HTN, Seizures, Hepatic impairment	More sensitive to CNS effects, Renal insufficiency, Headache, Constipation, N/V
<b>Liraglutide</b>	Angioedema, MEN-2, MTC	Constipation, Diarrhea, Hypoglycemia, Palpitations, N/V

Author Manuscript

Author Manuscript

Author Manuscript

Author Manuscript