Association between sarcopenic obesity and higher risk of type 2 diabetes in adults: A systematic review and meta-analysis

Dima Khadra, Leila Itani, Hana Tannir, Dima Kreidieh, Dana El Masri, Marwan El Ghoch

ORCID number: Dima Khadra (0000-0002-4711-6540); Leila Itani (0000-0003-3166-4945); Hana Tannir (0000-0002-3420-1965); Dima Kreidieh (0000-0001-5797-0460); Dana El Masri (0000-0001-9939-8116); Marwan El Ghoch (0000-0003-4277-4752).

Author contributions: El Ghoch M designed the work; Khadra D and El Ghoch M performed research, Itani L analyzed data, El Ghoch M, Itani L, Khadra D Tannir H, Kreidieh D, and El Masri D interpreted the data; El Ghoch M revised it critically for important intellectual content; Khadra D, Itani L, Tannir H, Kreidieh D, El Masri D and El Ghoch M drafted the work; all authors approved the final version of the manuscript.

Conflict-of-interest statement: All authors have no conflicts of interest to report.

PRISMA 2009 Checklist statement: The authors have read the PRISMA 2009 Checklist, and the manuscript was prepared and revised according to the PRISMA 2009 Checklist.

Abstract

BACKGROUND
The coexistence of sarcopenia and obesity is referred to as sarcopenic obesity (SO) and it has been hypothesized that the two components of SO may synergistically increase their negative effects. However, many uncertainties still surround this condition especially with regard to its potential negative effects on health outcomes.

AIM
To conduct a systematic review to determine the prevalence of sarcopenia among adults with overweight and obesity and to investigate whether SO was associated with a higher risk of type 2 diabetes (T2D).

METHODS
This study was conducted in adherence with the Preferred Reporting Items for Systematic Review and Meta-Analyses guidelines. Literature searches, study selection, methodology development and quality appraisal were performed independently by two authors and the data were collated by means of meta-analysis and narrative synthesis.

RESULTS
Of the 606 articles retrieved, 11 studies that comprised a total of 60118 adults with overweight and obesity of both genders met the inclusion criteria and were reviewed, revealing two main findings. First, the overall prevalence of sarcopenia is 43% in females and 42% in males who are with overweight and obesity. Second, the presence of SO increases the risk of T2D by 38% with respect to those without SO (OR = 1.38, 95%CI: 1.27-1.50).

CONCLUSION
A high prevalence of sarcopenia has been found among adults with overweight and obesity regardless of their gender and this condition seems to be associated
with a higher risk of T2D. Clinician should be aware of this scenario in their clinical practice for the better management of both obesity and T2D.

Key words: Obesity; Overweight; Sarcopenia; Type 2 diabetes; Reduced lean body mass

©The Author(s) 2019. Published by Baishideng Publishing Group Inc. All rights reserved.

Core tip: The coexistence of sarcopenia and obesity is referred to a phenotype termed sarcopenic obesity, defined as the increase in body fat deposition, and the reduction in lean mass and muscle strength. Since many uncertainties still surround this condition, especially with regard to its potential negative effects on health outcomes, we conducted this systematic review and found a high prevalence of sarcopenia among adults with obesity. Moreover, this condition seems to be associated with a higher risk of type 2 diabetes (T2D). Clinicians should be aware of this scenario in their clinical practice for better management of obesity and T2D.

Citation: Khadra D, Itani L, Tannir H, Kreidieh D, El Masri D, El Ghoch M. Association between sarcopenic obesity and higher risk of type 2 diabetes in adults: A systematic review and meta-analysis. World J Diabetes 2019; 10(5): 311-323
URL: https://www.wjgnet.com/1948-9358/full/v10/i5/311.htm
DOI: https://dx.doi.org/10.4239/wjd.v10.i5.311

INTRODUCTION

A condition that occurs because of the coexistence of sarcopenia and obesity has been termed sarcopenic obesity (SO)[1-7]. Many uncertainties still surround this phenomenon with regard to its definition and its potential negative effects on health outcomes, especially those related to obesity, namely the so-called cardio-metabolic diseases[8,9] such as type 2 diabetes (T2D), cardiovascular diseases, dyslipidaemia and metabolic syndrome[5,6,10-13]. In fact, it has been hypothesized that the two components of SO may synergistically increase their negative effects on health, however this is still a matter of debate[14-16].

Several studies have been conducted with a specific focus on determining the association between SO and T2D, however data regarding the contention that individuals with SO are likely to have poorer glycaemic profiles (i.e., hyperglycaemia, high HbA1c, insulin resistance, etc.) are still contradictory and require further clarification[11,17-23]. Moreover, to the best of our knowledge no systematic review posing this issue as a primary outcome has yet been conducted in order to provide an unbiased interpretation of the evidence published to date. In light of these considerations, we set out to systematically review the published literature with the aim of determining the prevalence of sarcopenia among adults with overweight and obesity and to investigate whether SO was associated with higher risk of T2D, in accordance with the PICO process[24-26] as detailed below: P – population: Individuals in the overweight or obese categories, however they were defined (i.e., body mass index (BMI), body fat percentage, waist circumference, etc.); I – seeking treatment (i.e., weight-loss or any other treatment if recruited from a clinical setting), otherwise non-treated if subjects were recruited from the general population; C – comparison: Comparison between individuals with sarcopenia and those without SO and with the healthy control group (when available); O – outcome: (i) Prevalence of SO however it was defined in the studies’ Methods section (i.e., low muscle mass, low muscle strength, low physical performance, increased visceral adiposity, increased waist circumference etc.,) and assessed (i.e., bioelectric impedance analysis (BIA), dual-energy X-ray absorptiometry (DXA), handgrip, etc,) among the entire obesity groups in the two genders; (ii) The prevalence of T2D however it was defined in the studies’ Methods section (i.e., fasting plasma glucose and glycated hemoglobin A1c, oral glucose tolerance test etc.,) in the SO and non-SO groups.

MATERIALS AND METHODS

The review conformed to the Preferred Reporting Items for Systematic Review and
Meta-Analyses guidelines\cite{28-30} and was registered in the PROSPERO Registry, No. CRD42018111931\cite{31}.

**Inclusion and exclusion criteria**
All studies that evaluated SO and T2D in adults were included, provided that they met the following criteria: (i) Studies written in English; (ii) Original research with a cross-sectional or longitudinal design; and (iii) Prospective or retrospective observational (analytical or descriptive), experimental or quasi-experimental controlled or non-controlled studies, documenting clearly the prevalence of SO, as well as the association or relationship between SO and T2D. No reviews or non-original articles (i.e., case reports, editorials, “Letters to the Editor” and book chapters) were included.

**Information source and search strategy**
The literature search was designed and performed independently in duplicate by two authors, namely the principal (DK) and the senior investigator (ME). The PubMed and Science direct databases\cite{32} were systematically screened using the MeSH terms and a manual search was carried out to retrieve other articles that had not been identified via the initial search strategy. Publication date was not considered an exclusion criterion for the purposes of this review.

**Study selection**
Two independent authors (DK and ME) screened the articles for their methodology and suitability for inclusion. The quality appraisal was conducted according to the Newcastle-Ottawa Scale (NOS), which relies on a 9-star system whereby scores of 0-3, 4-6 and 7-9 are considered poor, moderate and good quality, respectively\cite{33}. Consensus discussion was used to resolve disagreements between reviewers.

**Data collection process and data items**
The title and abstract of each paper were initially assessed by two independent authors (DK and ME) for language suitability and subject matter relevance, the selected studies were then assessed in terms of their suitability for inclusion and the quality of the methodology. The studies that passed both rounds of screening are presented in Table 1.

**Data synthesis**
The 11 studies that met the inclusion criteria have been presented as a narrative synthesis. In addition, a meta-analysis was conducted on the included studies using Med Calc. software\cite{34}. The Mantel Haenszel fixed and random effect models were used to estimate the overall effect size and 95%CI. The pooled estimate and 95%CI of the prevalence of SO among males and females in the included studies was estimated similarly.

**RESULTS**
The initial search retrieved 606 papers. After the first round of screening, 366 papers were excluded for: (i) Languages other than English; (ii) Non-human studies; and (iii) Dealing with obesity without sarcopenia, or the latter without the former. The second round of screening excluded 229 articles due to: (i) Inappropriate paper type, not original research articles (i.e., clinical reviews, Letters to the Editor, chapters in a book and case reports); (ii) Descriptions of SO, but not health-related outcomes; and (iii) An unclear definition of SO or identification of individuals with this condition. Accordingly, following the screening process, 11 articles were included in the systematic review and underwent narrative synthesis and meta-analysis (Figure 1). The NOS checklist proved that the studies were of a high quality (n = 11) (mean score = 7.36 points) (Table 2).

**Narrative synthesis**
In 2012, Sénéchal et al\cite{35} conducted a cross-sectional evaluation in which the authors assessed dynapenic obesity, defined as low leg muscle strength combined with abdominal obesity, in 1963 individuals with abdominal obesity. Of these patients, 566 had dynapenic obesity (data per gender is not available). Regardless of gender the mean age and mean BMI in the dynapenic obesity and non-dynapenic obesity groups were 65.4 ± 9.9 years and 29.9 ± 4.6 kg/m$^2$ and 65.5 ± 9.6 years and 30.8 ± 4.5 kg/m$^2$ respectively. Furthermore, 130 of the 566 individuals with dynapenic obesity had T2D compared to 196 of the 1397 individuals in the non-dynapenic obesity group.

One year later, Lu et al\cite{18} completed a cross-sectional study in which they assessed
| Study                  | Design            | Definition of SO                                                                 | Body composition       | Gender | Sample | Mean age | Mean BMI | Prevalence of Diabets | Prevalence of Sarcoprivic Obesity | Prevalence of Diabetes |
|-----------------------|-------------------|-------------------------------------------------------------------------------|------------------------|--------|--------|----------|---------|-----------------------|-------------------------------|-------------------------|
| Sénéchal et al[35], 2012 | Cross sectional   | Dynapenic obesity, defined as low leg muscle strength, combined with abdominal obesity | Kin-Corn dynamometer  | M-F    | T = 1963 Non DO: 65.5 ± 9.6; DO: 65.4 ± 8.9 | Non BMI: 29.9 ± 4.6; DO: 29.9 ± 4.6 | DO: n = 566/1963 (Did not distinguish in gender) | T2D: Non DO: n = 296; DO: n = 130 |
| Lu et al[18], 2013    | Cross sectional   | Defined by combination of total skeletal muscle mass/wt. (100) and BMI ≥ 25 kg/m² | BIA                    | M-F    | T = 180; M = 60; F = 120 | Non SO: 69.9 ± 7.3; SO: 61.1 ± 9.9 | Non BMI: 28.7 ± 2.6 | T2D: Non DO: n = 12/65; SO: n = 17/115 |
| Poggiogalle et al[36], 2015 | Cross sectional   | Defined by ASM/M² or ASM/BMI < 2SD of sex specific mean combined with assessment of FM and FFM | DXA                    | M-F    | T = 727; M = 141; F = 586 | 46.49 ± 13.73; 46.99 ± 13.76 | SO: n = 68/141 in males; n = 350/586 in females | T2D: Non DO: n = 36/71; SO: n = 34/71 |
| Ma et al[37], 2016    | Retrospective     | SO: BMI > 30 kg/m² and 24 h- UC < median                                      | DXA                    | M-F    | T = 310; M = 144; F = 166 | 71.8 ± 7.6 | 34.1 ± 4.0 | SO: n = 44/144 in males; n = 52/166 in females | T2D: Non DO: n = 105/700; SO: n = 165/855 |
| Xiao et al[38], 2017  | Retrospective     | FMI/FFMI ratio > 95 percentile of sex, BMI and ethnicity specific population-representative references | BIA                    | M-F    | T = 144; M = 45; F = 99 | Non SO: 56.6 ± 17.9; SO: 54.6 ± 10.1 | SO: ±7.6 | SO: ±8.3 | T2D: Non DO: n = 133/414; SO: n = 81/256 |
| Kang et al[39], 2017  | Cross sectional   | ASM/Wt < 1 SD the mean of the reference group, and BMI ≥ 25 kg/m²           | DXA                    | F      | T = 1555 | Non SO: 61.0 ± 4.4; SO: 62.91 ± 4.4 | Non BMI: 27.93 ± 0.11 | SO: n = 855/1555 (All females) | T2D: Non DO: n = 105/700; SO: n = 165/855 |
| Aubertin-Leheudre et al[40], 2017 | Cross sectional   | Dynapenic obesity, defined as low handgrip strength (≤ 19.9 in females; ≤ 31.9 in males), combined with BMI ≥ 25 kg/m² | Jamar Handheld Dynamometer | M-F    | T = 670; M = 213; F = 457 | Non SO: 76.3 ± 4.7; SO: 78.0 ± 4.6 | Non BMI: 34.9 ± 4.8 | SO: n = 77/213 in males; n = 179/457 in females | T2D: Non DO: n = 133/414; SO: n = 81/256 |
| Park et al[41], 2018  | Cross sectional   | SO defined by combination of SMI < 2 SD and WC ≥ 90 cm for men and ≥ 85 cm women | BIA                    | M-F    | T = 53818; M = 39820; F = 14998 | Non SO: 40.5 ± 9.2; SO: 40.5 ± 11.3 | SO: n = 3341; F = 3172 | SO: n = 391 |
| Kreidieh et al[42], 2018 | Cross sectional   | ALM/BMI < 0.512                                                              | BIA                    | F      | T = 154 | 33.26 ± 14.65 | 31.42 ± 4.94 | SO: n = 31 | T2D: Non DO: n = 3/123; SO: n = 4/31 |
SO defined as the coexistence of obesity (BMI ≥ 25 kg/m²) and sarcopenia based on the skeletal muscle index estimated by BIA. A sample of 180 individuals with obesity (60 males and 120 females) was recruited. Of the 60 males included in the sample 35 had SO compared to 80 of the 120 females. Regardless of gender the mean age and BMI in the SO group were 61.1 ± 9.9 years and 27.8 ± 2.6 kg/m² and 69.9 ± 7.3 years and 26.8 ± 1.6 kg/m² in the non-SO group. Moreover, 12 of the 65 patients in the SO group had T2D compared to 17 of the 115 patients in the non-SO group.

In early 2016, Poggiogalle et al. performed a cross-sectional study in which the authors assessed SO using DXA, with SO defined as the coexistence of obesity (BMI ≥ 30 kg/m²) and sarcopenia (ASM: height² < 6.54 and < 4.82 kg/m² for males and females respectively) or (ASM: weight < 0.2827 and < 0.2347 for males and females respectively). This study enrolled a sample of 727 individuals with obesity (141 males and 586 females), with mean ages of 45.63 ± 13.53 and 45.76 ± 13.58 years, and mean BMIs of 37.56 ± 5.99 and 37.80 ± 5.77 kg/m² respectively for each gender. Of the 141 male patients 68 had SO, while 350 of the 586 females had the condition. In addition, 155 of the 418 patients had pre-diabetes or T2D in the SO group compared to 70 of the 309 patients in the non-SO group.

In the same year, Ma et al. performed a cross-sectional evaluation on SO defined by BMI and sex-specific 24-h urinary creatinine excretion, in 310 patients (166 females and 144 males) with obesity (BMI ≥ 30 kg/m²). Fifty-four of the 144 males and 52 of the 166 females had SO. The mean BMI and age of the SO group were 34.1 ± 4.0 kg/m² and 71.8 ± 7.6 years, while they were 34.9 ± 4.4 kg/m² and 67.8 ± 6.8 years in the non-SO group. Furthermore, 40 of the 106 patients had T2D in the SO group in comparison to 51 of the 204 patients in the non-SO group.

In 2017, Xiao et al. performed a retrospective study on the prevalence of SO and its association with health outcomes in patients seeking weight loss treatment in a bariatric surgery setting. Body composition analysis was conducted by means of BIA and SO was defined by a fat mass/fat-free mass index (FMI: FFMI) ratio greater than the 95th percentile of sex, BMI and ethnicity-specific population-representative references. A sample of 144 adults with obesity (99 females and 45 males) were enrolled, with a mean age of 55.6 ± 11.5 years and a mean BMI of 46.6 ± 8.4 kg/m². Of the 144 patients included in the sample 73 had SO (data per gender is not available). The mean age and BMI of the SO group were 44.0 ± 7.6 kg/m², and 67.8 ± 6.8 years in the non-SO group, respectively. Furthermore, 34 of the 73 patients had T2D in the SO group in comparison to 51 of the 204 patients in the non-SO group.

In 2017, Kang et al. conducted a large cross-sectional study to assess the association between SO and metabolic syndrome in postmenopausal women. SO was defined by the co-existence of sarcopenia (ASM: weight < 1 standard deviation below the mean of the reference group) and a BMI cut-off point for obesity which referred to a score of 25 kg/m² on the basis of the Asia-Pacific obesity criterion. The study included 1555 females with obesity, of whom 855 had SO, with a mean age of 62.91 ± 4.4 years and a mean BMI of 27.93 ± 0.11 kg/m². On the other hand, 700 did not have SO and had a mean age of 61.05 ± 0.44 years and a mean BMI of 26.80 ± 0.07 kg/m².

**Table 1:**

| **Cross sectional** | **ALM/BMI < 0.789, (ALM/WT) × 100% < 25.72, and (ALM/WT) × 100% < 29.60** | **BIA** | **Handgrip strength Gait speed** | **DXA** | **M** | **T = 72** | **T = 525** | **Non SO: 75.9 ± 4.7; SO: 80.3 ± 6.5** | **Non SO: 30.7 ± 3.4; SO: 27.2 ± 2.3** | **n = 80** | **High fasting glucose or diabetes medications: Non SO: n = 177/445; SO: n = 29/80** |
|---------------------|---------------------------------------------------------------------------------|--------|---------------------------------|---------|--------|------------|------------|-------------------------------------|-------------------------------|----------|-----------------------------------|
| **Scott et al.**     | **Cross sectional (includes a longitudinal part)**                              | **M**  | **T = 72**                      |         | **M**  | **T = 525** | **Non SO: 75.9 ± 4.7; SO: 80.3 ± 6.5** | **Non SO: 30.7 ± 3.4; SO: 27.2 ± 2.3** | **n = 80** | **High fasting glucose or diabetes medications: Non SO: n = 177/445; SO: n = 29/80** |

SO: Sarcopenic obesity; DO: Dynapenic obesity; BMI: Body mass index; M: Male; F: Female; BIA: Bioelectric impedance analysis; T2D: Type 2 diabetes; DXA: Dual-energy X-ray absorptiometry.
addition, 165 of the 855 patients had T2D in the SO group while 105 of the 700 patients in the non-SO group had T2D.

In the same year, a cross-sectional study by Aubertin-Leheudre et al[40] aimed to examine the association between dynapenic obesity and metabolic risk factors in older adults (age ≥ 70 years). Dynapenic obesity was defined as low handgrip strength (≥ 19.9 in females; ≤ 31.9 in males) combined with a BMI of ≥ 30 kg/m². The study included 670 participants with obesity (213 males and 457 females), of whom 256 had dynapenic obesity, with a mean age of 78.0 ± 4.6 years and a mean BMI of 34.9 ± 4.8 kg/m², and 414 did not have dynapenic obesity, with a mean age of 76.3 ± 4.7 years and a mean BMI of 35.6 ± 4.8 kg/m². Furthermore, 81 of the 256 individuals in the dynapenic obesity group had T2D while 133 of 414 individuals in the non-dynapenic obesity group had T2D.

In 2018, Park et al[41] conducted a large cross sectional study in two sites, which included a total of 53818 adults with overweight and obesity of both genders (38820 males and 14998 females), of whom 6513 had SO defined as below two standard deviations of the mean of the skeletal muscle mass index for young adults assessed by BIA and a waist circumference of ≥ 90 cm for men and ≥ 85 cm for women. The mean age and BMI of the individuals with obesity only were 40.5 ± 9.2 years and 26.9 ± 2.2 kg/m² compared to those with SO who had a mean age of 40.0 ± 11.3 years and a mean BMI of 30.7 ± 3.4 kg/m². Moreover, 391 of the 6513 patients had T2D in the SO group compared to 2176 of the 47305 patients in the non-SO group.

In 2018, Kreidieh et al[42] conducted a cross sectional controlled study in which body composition measurements were conducted by BIA using a definition that in addition to appendicular lean mass (ALM) also involved BMI, and patients were considered affected by SO if ALM:BMI < 0.512. The study included 154 females with overweight and obesity with a mean age of 33.26 ± 14.65 years and a mean BMI of 31.42 ± 4.94 kg/m². Of the 154 female patients 31 had SO. Moreover, four of the 31 patients had T2D in the SO group compared to three of the 123 patients in the non-SO group.

In 2018, Khazem et al[43] performed a cross-sectional controlled study on 72 adult
Table 2 Quality assessment of the included studies

| Author                     | Selection | Represents cases with independent validation | Cases are consecutive or obviously representative | Controls from the community | Controls have no history of sarcopenia | Comorbidities | Controls are comparable for the most important factors | Control for any additional factor | Ascertainment of exposure | Total score |
|----------------------------|-----------|-----------------------------------------------|-----------------------------------------------------|----------------------------|---------------------------------------|---------------|--------------------------------------------------------|-------------------------------|----------------------------|-------------|
| Sénéchal et al[35], 2012   | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Lu et al[18], 2013         | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Poggiogalle et al[36], 2016| 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Ma et al[41], 2017         | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Xiao et al[38], 2017       | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Kang et al[40], 2017       | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Aubertin-Leheudre et al[42], 2017 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 7 |
| Park et al[43], 2018       | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Scott et al[44], 2018      | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Kreidieh et al[45], 2018   | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |
| Khazem et al[46], 2018     | 1         | 1                                             | 1                                                   | 1                          | 1                                     | 1             | 1                                                     | 1                             | 1                          | 7           |

Newcastle-Ottawa Scale (NOS) for longitudinal and cross-sectional studies. Yes = 1, No (not reported, not available) = 0; Studies with scores of 0-3, 4-6, 7-9 were considered as low, moderate and high quality, respectively.

males with overweight and obesity with a mean age of 32.79 ± 13.65 years and a mean BMI of 33.69 ± 5.84 kg/m². In this study the authors used three different definitions proposed by Batsis et al[44], Levine and Crimmins[21], and Oh et al[45] based on ALM:BMI.
and (ALM: weight) × 100% to define SO. Body composition was assessed by BIA. Based on each formula the prevalence of SO varied between 23.9% and 69.4%. However, based on the definition that was revealed to be more useful from the clinical perspective, 50 of the 72 patients had a reduced lean body mass with a prevalence of 69.4%. Moreover, three of the 50 patients had T2D in the SO group in comparison to one of the 22 patients in the non-SO group.

Finally, in 2018 Scott et al.[44] conducted a large sampled study that aimed to investigate the cross-sectional association between SO and components of metabolic syndrome in community-dwelling older men. SO was defined by the co-existence of sarcopenia as ALM/height < 7.26 kg/m² combined with handgrip strength < 30 kg and/or low gait speed ≤ 0.8 m/s, while obesity was defined as a body fat percentage of ≥ 30%. The study included 525 males with obesity, of whom 80 had SO, with a mean age of 80.3 ± 6.5 years and mean BMI of 27.2 ± 2.3 kg/m² and 445 did not have SO, with a mean age of 75.9 ± 4.7 years and mean BMI of 30.7 ± 3.4 kg/m². Furthermore, 29 of the 80 individuals in the SO group had T2D in comparison to 177 of the 445 individuals in the non-SO group.

Meta-analysis
The meta-analysis estimated the overall prevalence of SO among males and females. With high heterogeneity among the included studies, a random effect model was considered for the estimation of the overall prevalence of SO. The forest plots in Figures 2 and 3 show that SO affected 43% (95% CI: 28-59) of females and 42% (95% CI: 31-53) of males. In addition, the overall odds ratios of T2D in patients with SO as compared to those without SO are presented in Figure 4. The fixed effect weighted pooled odds for T2D in patients with SO indicated an increased risk of T2D of approximately 38% compared to those without SO (OR: 1.38, 95% CI: 1.27-1.50). The heterogeneity analysis revealed moderate variability (I² = 60%).

DISCUSSION
This systematic review aimed to provide benchmark data on the prevalence of sarcopenia in individuals with overweight and obesity and to assess any potential association between SO and T2D in this population. The major finding is that sarcopenia seems to affect approximately 40%-45% of individuals with overweight and obesity of both genders, and the co-existence of both conditions, namely sarcopenia and excess weight/obesity increases the risk of T2D by nearly 38% when compared with those who had excess weight or obesity alone. The underlying mechanism behind this association is still unclear, however it seems that there is a bidirectional interaction between obesity, chronic inflammation, insulin resistance and sarcopenia[19]. In fact, the chronic inflammation plays an important role in the pathogenesis of T2D. For this reason we speculate that coexistence of both obesity and sarcopenia under the so-called phenotype “SO”, may have a synergistic effect with chronic inflammation being a common “denominator” seen in both conditions, which seems to exacerbate further glucose metabolism impairment (i.e., insulin resistance, pre-diabetes and T2D)[19].

The clinical implication of this review’s findings is the awareness of the high prevalence of sarcopenia in the overweight/obese population that should be raised among clinicians and patients. Secondly, these results reveal the importance of screening for SO in individuals affected by excess weight and obesity, since this condition also seems to be strongly associated with T2D.

This systematic review has certain strengths. To the best of our knowledge this is the first systematic review to assess the overall prevalence of SO in males and females with overweight and obesity. In fact, the studies that have been conducted on this topic reported varying levels of prevalence that ranged between 0 and 100%, depending on the applied definition of SO[30,46]. Higher prevalence tends to be reported in studies that accounted for body mass (i.e., BMI), whereas a lower prevalence is reported in those that did not[10,46]. A low prevalence may also be explained by the use of definitions that have primarily been developed from studies in older cohorts and these may not be applicable to younger adults[46].

However, this systematic review also has certain limitations. Foremost, our results need to be interpreted with caution with regard to the association between SO and the prevalence of T2D, since the cross-sectional design of the studies (i.e., non cohort), included in our systematic review indicates only simple associations between SO and T2D at best and does not provide solid information regarding any causal relationships between the two conditions[30,31]. In other words, these studies lack evidence to determine if SO may lead to the onset or deterioration of T2D, since very few studies...
have longitudinally investigated the “real” effects of SO on health\cite{52}. These shortcomings in the current research indicate the need to design longitudinal studies to clarify the real effect of SO on the onset and progression of T2D.

In conclusion, a high prevalence of sarcopenia has been found among adults with overweight and obesity regardless of their gender, and this condition seems to be associated with a higher risk of T2D. Clinicians should be aware of this scenario in their clinical practice for better management of both obesity and T2D.
ARTICLE HIGHLIGHTS

Research background
The coexistence of sarcopenia and obesity has been termed as sarcopenic obesity (SO). Several studies have been conducted in order to determine any potential association between SO phenotype and type 2 diabetes (T2D). However, the available data are still contradictory and require further clarification.

Research motivation
To our knowledge no systematic review on the primary outcome related to the association between SO and T2D has been conducted yet to provide an unbiased interpretation of the evidence published to date.

Research objectives
We set out to systematically review the published literature with the aim of determining the prevalence of sarcopenia among adults with overweight and obesity and to investigate whether SO was associated with higher risk of T2D.

Research methods
The review conformed to the Preferred Reporting Items for Systematic Review and Meta-Analyses guidelines, and data were collated by means of narrative synthesis and meta-analysis.

Research results
The prevalence of SO in adult with overweight and obesity is 43% in females and 42% in males, and the presence of this condition increases the risk of T2D by 38% with respect to those without SO.

Research conclusions
A high prevalence of sarcopenia has been found among adults with overweight and obesity regardless of their gender, and this condition seems to be associated with a higher risk of T2D. The clinical implication of our findings is to raise awareness of the high prevalence of this phenotype in the overweight/obese population, and the importance of screening for SO in...
individuals affected by excess weight, since this condition seems to be strongly associated with T2D. However, our results need to be interpreted with caution with regard to the association between SO and the prevalence of T2D, since the cross-sectional design of the studies included in our systematic review indicates only associations between the two conditions and that does not provide information regard the causal relationships.

**Research perspectives**

The current research indicates the need to design longitudinal studies to clarify the real effect of SO on the onset and progression of T2D. In other words, the available studies lack in evidence to determine if SO may lead to the onset or deterioration of T2D, since very few studies have longitudinally investigated the “real” effects of SO on health.

**REFERENCES**

1. Choi KM. Sarcopenia and sarcopenic obesity. *Korean J Intern Med* 2016; 31: 1054-1060 [PMID: [227809450 DOI: 10.3904/kim.2016.193]
2. Polyzos SA, Margioris AN. Sarcopenic obesity. *Hormones (Athens)* 2018; 17: 321-331 [PMID: [30014320 DOI: 10.1007/s42001-018-0049-s-n]
3. Bouchonville MF, Villareal DT. Sarcopenic obesity: how do we treat it? *Curr Opin Endocrinol Diabetes Obes* 2013; 20: 412-419 [PMID: [23974769 DOI: 10.1097/01.med.0000437071.14867.7]
4. Choi KM. Sarcopenia and sarcopenic obesity. *Endocrinol Metab (Seoul)* 2013; 28: 86-89 [PMID: [23966569 DOI: 10.3803/EmM.2013.28.2.86]
5. dos Santos EP, Gadilha AB, Safons MP, Nobrega OT, Oliveira RJ, Lima RM. Sarcopenia and sarcopenic obesity classifications and cardiometabolic risks in older women. *Arch Gerontol Geriatr* 2014; 59: 56-61 [PMID: [24769935 DOI: 10.1016/j.archger.2014.03.012]
6. Lim S, Kim JH, Yoon JW, Kang SM, Choi SH, Park YJ, Kim KW, Lim JY, Park KS, Jang HC. Sarcopenic obesity: prevalence and association with metabolic syndrome in the Korean Longitudinal Study on Health and Aging (KLONSHA). *Diabetes Care* 2010; 33: 1652-1654 [PMID: [20460442 DOI: 10.2337/dc10-0107]
7. Kwong B, Lim JY, Lee J, Choi NK, Ahn YO, Park BJ. Prevalence rate and associated factors of sarcopenic obesity in Korean elderly population. *J Korean Med Sci* 2012; 27: 748-755 [PMID: [227787639 DOI: 10.3346/jkms.2012.27.7.748]
8. Visser M, van Wouwe PM, Vink H, van Leeuwen PA, de Mol BA. Sarcopenic obesity is associated with adverse clinical outcome after cardiac surgery. *Nutr Metab Cardiovasc Dis* 2013; 23: 511-518 [PMID: [23517879 DOI: 10.1016/j.numecd.2013.12.001]
9. Park SH, Park JH, Song PS, Kim DK, Kim KH, Seol SH, Kim HK, Jang JH, Lee JG, Park HY, Park J, Shin KJ, Kim DI, Moon YS. Sarcopenic obesity as an independent risk factor of hypoglycemia. *J Am Soc Hypertens* 2013; 7: 420-425 [PMID: [23910010 DOI: 10.1016/j.jash.2013.06.002]
10. Steinholm S, Harris TB, Rantanen T, Visser M, Kritchevsky SB, Ferrucci L. Sarcopenia definition, cause and consequences. *Curr Opin Clin Nutr Metab Care* 2008; 11: 693-700 [PMID: [18827572 DOI: 10.1097/MCO.0b013e32833c377d]
11. El Ghoch M, Calugi S, Dalle Grave R. Sarcopenic Obesity: Definition, Health Consequences and Clinical Management. *The Open Nutrition Journal* 2018; 12: 70-73 [DOI: 10.2174/18742881201006070]
12. Baek SJ, Nam GE, Han KD, Choi SW, Jung SW, Bok AR, Kim YH, Lee KS, Han BD, Kim DH. Sarcopenia and sarcopenic obesity and their associations with dyslipidemia in Korean elderly men: the 2008-2010 Korea National Health and Nutrition Examination Survey. *J Endocrinol Invest* 2014; 37: 247-260 [PMID: [24615361 DOI: 10.4086/0161-013-0011-3]
13. Messier V, Karelis AD, Lavoie ME, Brochu M, Faraj M, Strychar I, Rabasa-Lhoret R. Metabolic profile and quality of life in class 1 sarcopenic overweight and obese postmenopausal women: a MONET study. *Appl Physiol Nutr Metab* 2009; 34: 18-24 [PMID: [19234585 DOI: 10.1113/jcn.2008.273059]
14. Gusmao-Sena MH, Curvello-Silva K, Barreto-Medeiros JM, Da-Cunha-Daltro CH. Association between sarcopenic obesity and cardiovascular risk: where are we? *Nutr Hosp* 2016; 33: 592 [PMID: [27759996 DOI: 10.20960/nh.592]
15. Barazzoni R, Bischoff SC, Boirie Y, Busetto L, Cederholm T, Dicker D, Toplak H, Van Gossum A, Yannuk V, Vetter R. Sarcopenic obesity: Time to meet the challenge. *Clin Nutr* 2018; 37: 1787-1793 [PMID: [29857921 DOI: 10.1016/j.clnut.2018.04.018]
16. Barazzoni R, Bischoff SC, Busey L, Cederholm T, Dicker D, Toplak H, Van Gossum A, Yannuk V, Vetter R. Sarcopenic Obesity: Time to Meet the Challenge. *Obes Facts* 2018; 11: 294-305 [PMID: [23016792 DOI: 10.1159/000490261]
17. Kim TN, Park MS, Lim KJ, Choi HY. Yang SJ, Yoo HJ, Kang HJ, Song W, Choi H, Baik SH, Choi DS, Choi K. Sarcopenia and sarcopenic obesity and insulin resistance, inflammation, and vitamin D status: the Korean Sarcopenic Obesity Study. *Clin Endocrinol (Oxf)* 2013; 78: 525-532 [PMID: [22563924 DOI: 10.1111/j.1365-2265.2012.04433.x]
18. Lu CW, Yang KC, Chang JH, Lee JT, Chen CY, Huang KC. Sarcopenic obesity is closely associated with metabolic syndrome. *Obes Res Clin Pract* 2013; 7: e301-e307 [PMID: [23406159 DOI: 10.1016/j.orec.2012.02.001]
19. Srikantan P, Heveren AL, Karlamangla AS. Sarcopenia exacerbates obesity-associated insulin resistance and dysglycemia: findings from the National Health and Nutrition Examination Survey III. *PLoS One* 2010; 5: e10805 [PMID: [22421977 DOI: 10.1371/journal.pone.0010805]
20. Murai J, Nishizawa H, Otsuka A, Fukuda S, Tanaka Y, Nagao H, Sakai Y, Suzuki M, Yokota S, Tada H, Doi M, Fujisawa Y, Kita S, Funahashi T, Maeda N, Nakamura T, Shimomura I. Low muscle quality in Japanese type 2 diabetic patients with visceral fat accumulation. *Cardiovasc Diabetol* 2018; 17: 112 [PMID: [30077185 DOI: 10.1186/s12933-018-0755-z]
21. Levine ME, Crimmins EM. The impact of insulin resistance and inflammation on the association between sarcopenic obesity and physical functioning. *Obesity (Silver Spring)* 2012; 20: 2101-2106 [PMID: [22310233 DOI: 10.1038/oby.2012.20]
22. Kim K, Park SM. Association of muscle mass and fat mass with insulin resistance and the prevalence of metabolic syndrome in Korean adults: a cross-sectional study. *Sci Rep* 2018; 8: 2703 [PMID: [29246839]
Sarcopenic Obesity in Adults with Class II/III Obesity Using Different Diagnostic Criteria.

J Nutr Metab, Sharma AM, Forhan M, Siervo M, Padwal RS, Prado CM. Prevalence of sarcopenia in older men: The Concord Health and Ageing in Men Project.

Exp Gerontol, Cumming R, Naganathan V, Blyth F, Le Couteur DG, Handelsman DJ, Seibel M, Waite LM, 2015; 108, 105-115. but elderly women had a higher prevalence of sarcopenic obesity in a population of Korean adults.

J Nutr, Manini TM, Fielding R, Newman A, Church T, 2017; 9-14. National Health and Nutritional Examination Surveys from 2008 to 2011.

J Bone Metab, Do JG, Lee YT, Yoon KJ. Sarcopenic obesity associated with high-sensitivity C-reactive protein in age and sex comparison: a two-center study in South Korea.

J Am Med Dir Assoc, Cohen JF, Deeks JJ, Gatsonis C, Hooft L, Hunt HA, Hyde CJ, Korevaar DA, Leeflang MMG, Macaskill P, Kleijnen J, Moher D. The PRISMA statement for reporting health care interventions: explanation and elaboration. Br J Sports Med.

Clinical Nutr, Aubertin-Leheudre M, Itani L, Kreidieh D, El Ghoch M. Reduced Lean Body Mass and Cardiometabolic Diseases in Adult Males with Obesity and Obesity: A Pilot Study. Int J Environ Res Public Health, 2018; 15 [PMID: [30563167 DOI: 10.3390/ijerph15127754]

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 2015; 35: 1031-1039 [PMID: [26472145 DOI: 10.1016/j.nutres.2015.08.006]

Oh C, Jho S, Kim JS, Kim HS. Body composition changes were related to nutrient intakes in elderly men and elderly women had a higher prevalence of sarcopenic obesity in a population of Korean adults.

Nutr Res 2015; 35: 1-6 [PMID: [25524331 DOI: 10.1016/j.nutres.2014.07.018]

Scott D, Cumming R, Naganathan V, Blyth F, Le Couture DG, Handelsman DJ, Seibel M, Waite LM, Hirani V. Associations of sarcopenia obesity with the metabolic syndrome and insulin resistance over five years in older men: The Concord Health and Ageing in Men Project. Exp Gerontol 2018; 108: 99-105 [PMID: [29649572 DOI: 10.1016/j.exger.2018.04.006]

Johnson Stoklosa CA, Sharma AM, Forhan M, Siervo M, Padwal RS, Prado CM. Prevalence of Sarcopenic Obesity in Adults with Class II/III Obesity Using Different Diagnostic Criteria. J Nutr Metab 2017; 2017: 7387618 [PMID: [2942144 DOI: 10.1155/2017/7387618]
Lee DC, Shook RP, Drenowatz C, Blair SN. Physical activity and sarcopenic obesity: definition, assessment, prevalence and mechanism. *Future Sci OA* 2016; 2: FSO127 [PMID: [28031974 DOI: 10.4155/fsoa-2016-0028]]

El Ghoch M, Rossi AP, Calugi S, Rubele S, Soave F, Zamboni M, Chignola E, Mazzali G, Bazzani PV, Dalle Grave R. Physical performance measures in screening for reduced lean body mass in adult females with obesity. *Nutr Metab Cardiovasc Dis* 2018; 28: 917-921 [PMID: [30017438 DOI: 10.1016/j.numecd.2018.06.008]]

Mahajan A. Limitations of cross-sectional studies. *Neurol India* 2015; 63: 1006-1007 [PMID: [26588658 DOI: 10.4103/0028-3886.170108]]

Setia MS. Methodology Series Module 3: Cross-sectional Studies. *Indian J Dermatol* 2016; 61: 261-264 [PMID: [27293245 DOI: 10.4103/0019-5154.182416]]

Cuthbertson DJ, Bell JA, Ng SY, Kemp GJ, Kivimaki M, Hamer M. Dynapenic obesity and the risk of incident Type 2 diabetes: the English Longitudinal Study of Ageing. *Diabet Med* 2016; 33: 1052-1059 [PMID: [26479063 DOI: 10.1111/dme.12991]]
