# Relationship between Low Muscle Mass and Metabolic Syndrome in Elderly People with Normal Body Mass Index 

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Background: Metabolically obese normal weight (Wt) occurs in people who have high prevalence of cardiovascular disease even with normal body mass index (BMI). In this study, we determined the relationship between low muscle mass and metabolic syndrome which is known to be associated with cardiovascular disease in aged people with normal BMI. Methods: This study used the data of Korea National Health and Nutrition Examination Survey. Aged people ( $\geq 65$ years) with normal BMI ( 18.5 to $22.9 \mathrm{~kg} / \mathrm{m}^{2}$ ) were enrolled. Low muscle mass class I was defined as appendicular skeletal muscle divided by Wt 1 to 2 standard deviation below the mean of gender specific young healthy adult and class II was defined as below 2 standard deviations. Results: The prevalence of low muscle mass class I and II were $18.8 \%$ and $5.2 \%$ in men and $11.2 \%$ and $2.2 \%$ in women, respectively. In men and women, low muscle mass and metabolic syndrome was associated with each other after adjusting for age. After further adjusting for BMI, smoking status, alcohol consumption, and exercise frequency, the relationship was maintained in aged men. Moreover in aged men, the risk of metabolic syndrome was increased according to the grade increase in low muscle mass class. Conclusions: In the aged with normal BMI, low muscle mass was a risk factor of metabolic syndrome. Furthermore, in normal BMI men, when the degree of low muscle mass was enhanced, the risk of metabolic syndrome was increased.

Key Words: Aged, Body mass index, Metabolic syndrome, Muscle mass

## INTRODUCTION

Body mass index (BMI) is an indicator of obesity that is the most widely used as it is easy and convenient to measure and calculate. However, there are limits of using BMI only to assess obesity, because when a case has a large body fat mass but appropriate overall weight (Wt) is classified as normal. On the contrary, any case with a small fat percentage can be classified as obese if having more muscle mass.[1] Therefore there are limits in predicting the risks of cardiovascular diseases and other metabolic diseases by defining obesity only using a simple BMI. In order to supplement those limits, in recent days, those with normal BMI but high-
er morbidity to metabolic diseases are defined as metabolically obese normal-Wt (MONW).[2]

As getting old, the body fat mass is increased and appendicular skeletal muscle mass (ASM) is reduced even if there is no change in the Wt.[3] Such loss of muscle mass and decreasing of strength is defined as sarcopenia.[4] When the muscles are reduced, it becomes frailty and its function is deteriorated,[5] so, it is known to ultimately increase mortality.[6] When reviewed studies conducted so far, there are some cross-sectional studies reported that the prevalence of metabolic syndrome and cardiovascular diseases has been increased in sarcopenia,[7] and some studies presented that the reduction of muscle mass did not have any independent associations with metabolic syndrome in the aged people.[8] Sarcopenia and the prevalence of metabolic syndrome had shown a large difference depending on various ways of defining sarcopenia, and according to a study, it had reported when defined sarcopenia by using the ASM adjusted for Wt, the level of insulin resistance and cholesterol was displayed as high in the group with sarcopenia, but when defined sarcopenia by using the value adjusted for height ( Ht ), there was no association presented.[9] Like this, the association of muscle mass with the risk of metabolic diseases is still controversial.

When BMI is normal, there is a possibility that obesity is not diagnosed in primary care so that it is difficult to control lifestyle such as diet and exercise, and is likely to have delayed diagnosis and treatment of chronic diseases. Therefore, studies analyzing the groups vulnerable to the chronic diseases despite normal Wt are required, but there are no studies so far, performed in Korean population to analyze the association between the metabolic syndrome and the presence/absence of sarcopenia. To this end, this study aimed to identify how skeletal muscle mass adjusted for Wt influence on the metabolic syndrome in those aged $\geq 65$ years with normal BMI, based on the Korea National Health and Nutrition Examination Survey (KNHANES) that has representativeness in Korean population.

## METHODS

## 1. Study subjects

The KNHANES was started in 1998, and currently it is conducted by the Korea Centers for Disease Control and Pre-
vention (KCDC). It had introduced a rolling sampling survey so that it can represent the entire population of Korea. This nationally representative survey is composed with three component surveys: health interview, health examination including anthropometric measures and clinical profiles with three component surveys: health interview, health examination and nutrition survey to identify the actual status of nutrition. The KNHANES had measured bone density, muscle mass and body fat mass using a dual energy X-ray absorptiometry (DXA; Hologic Inc., Bedford, MA, USA) from July, 2008 to May, 2011. This study was conducted in adult men and women aged $\geq 65$ years who are applicable to $\mathrm{BMI} \geq 18.5 \mathrm{~kg} / \mathrm{m}^{2}$ and $<23.0 \mathrm{~kg} / \mathrm{m}^{2}$ which is the Asia-Pacific Region Criteria of World Health Organization (WHO) among those who had completed DXA measurement within applicable period. Among the study population, patients with myocardial infarction, angina pectoris, stroke, renal failure, liver cirrhosis, or malignant tumor were excluded from analysis, so that a total of 674 men and 733 women were included in the analysis. All of the study participants were fully briefed about the study, and submitted informed consents for participation in health interview, health examination and nutrition survey.

## 2. Study method

The skeletal muscle mass was evaluated using the value (ASM/Wt, \%) obtained by dividing ASM measured through DXA modified the definition of sarcopenia presented by Jassen, by Wt.[10,11] Among adults aged between 20 to 39 years old who had participated in the KNHANES of the same period, the healthy men and women were set as the reference group, except patients with malignant tumor, stroke, myocardial infarction, angina pectoris, liver cirrhosis, heart failure, diabetes, hypertension, tuberculosis or asthma. Low muscle mass class 1 was defined as ASM divided by Wt 1 to 2 standard deviation below the mean ASM /Wt (\%) value of gender specific young healthy adult and class 2 was defined as below 2 standard deviations.[9,12] According to the defining method of this study, those men with ASM/Wt (\%) $\geq 30.52 \%$ were classified as normal group, whereas those men with ASM/Wt (\%) $\geq 27.85 \%$ and less than $30.52 \%$ were classified as low muscle mass class 1, and with ASM/Wt (\%) less than $27.85 \%$ were classified as low muscle mass class 2 . In women, they were classified as normal and low muscle mass class 1 and 2 , having $23.80 \%$ and

### 21.35\% as references.

Among a variety of diagnostic criteria for metabolic syndrome, this study had adapted the National Cholesterol Education Program Adult Treatment Panel III (Modified NCEP ATP III) Criteria revised in 2004, and applied the Waist Circumference cut-off points for defining abdominal obesity in Koreans. When satisfied three or more of the following 5 criteria, the case was defined as metabolic syndrome.[13,14] 1) Abdominal obesity: the waist circumference $\geq 90 \mathrm{~cm}$ for men, $\geq 85 \mathrm{~cm}$ for women respectively, 2) Hypertension: systolic blood pressure (BP) $\geq 130 \mathrm{mmHg}$ or diastolic BP $\geq 85 \mathrm{mmHg}$ or taking anti-hypertensive medication, 3) Hypertriglyceridemia: the level of triglyceride $\geq 150 \mathrm{mg} /$ dL or taking medication for dyslipidemia treatment, 4) Low high-density lipoproteins (HDL) cholesterolemia: HDL cholesterol less than $40 \mathrm{mg} / \mathrm{dL}$ for men, less than $50 \mathrm{mg} / \mathrm{dL}$ for women, or taking medication for dyslipidemia treatment, 5) Diabetes: fasting blood glucose $\geq 100 \mathrm{mg} / \mathrm{dL}$, taking hypoglycemic agent or insulin therapy.

## 3. Statistics

Taking into account the Wt per investigation area and the number of subjects for each test item per year, suggested by the KNHANES according to the test items, Integrated Wt was generated and used in the study analysis. Continuous variables had presented the calculated values using the mean $\pm$ standard deviation (SD), whereas categorical variables had presented them using the estimated percentage. Through multiple logistic regression, variables that were likely to affect the metabolic syndrome were calibrated and the association of the ASM adjusted for Wt with the metabolic syndrome in normal Wts.

The study had used SPSS version 20.0 (SPSS Inc., Chicago, IL, USA) software, if the $P$-value ( $P$ value, $P$ ) and the trend value $P$ ( $P$ for trend) of less than 0.05 was considered statistically significant.

## RESULTS

Out of 674 male participants in the study, it was presented that $18.8 \%$ was low muscle mass class 1 and $5.2 \%$ was low muscle mass class 2 . Among 733 female participants, 11.2\% and 2.2\% were classified as low muscle mass class 1 and 2 respectively.

In men, the mean age of normal group was 71.8 years
old, low muscle mass class 1 was 73.3 years old, and low muscle mass class 2 was 75.0 years old, indicating when higher the low muscle mass class got, older the mean age was ( $P=0.001$ ). There were differences in BMI between three groups, presenting $20.6 \mathrm{~kg} / \mathrm{m}^{2}, 21.4 \mathrm{~kg} / \mathrm{m}^{2}$ and $21.6 \mathrm{~kg} / \mathrm{m}^{2}$ in each group respectively ( $P<0.001$ ). There were differences also in the waist circumferences between three groups, presenting 77.3 cm in men with normal skeletal muscle mass, whereas it was 81.2 cm in low muscle mass class 1 , and 83.5 cm in low muscle mass class 2, and there were significant differences between three groups ( $P<0.001$ ). Systolic BP was statistically significantly higher in the group with less skeletal muscle mass compared to the group with normal skeletal muscle mass ( $P=0.024$ ), and when compared the group with low muscle mass class 2 to other groups, there were also significant differences presented ( $P=0.007$ ). Fasting blood glucose in three groups were $99.4 \mathrm{mg} / \mathrm{dL}$, $111.1 \mathrm{mg} / \mathrm{dL}$ and $104.3 \mathrm{mg} / \mathrm{dL}$ respectively, presenting there were differences between the group with less skeletal muscle mass and the normal group ( $P<0.001$ ). The level of triglyceride had presented significant differences between ( $P=0.006$ ), but the level of HDL cholesterol had presented statistically significant differences when compared the group with low muscle mass class 2 to other groups ( $P=0.028$ ) (Table 1).

In women, there was no significant correlation presented from the skeletal muscle mass and age. The waist circumference was 74.5 cm in the normal group, while according to the levels of skeletal muscle mass, low muscle mass class 1 and 2 had presented 79.2 cm and 82.0 cm respectively, showing that low muscle mass class 2 presented the highest value ( $P<0.001$ ). The levels of triglyceride in three groups were $137.1 \mathrm{mg} / \mathrm{dL}, 154.6 \mathrm{mg} / \mathrm{dL}$ and $177.7 \mathrm{mg} / \mathrm{dL}$ respectively, presenting there were differences between three groups ( $P<0.05$ ), however, the items such as BP, fasting blood glucose and HDL cholesterol had not presented any statistically significant differences between three groups ( $P>0.05$ ) (Table 2).
In men, as progressing from the normal, low muscle mass class 1 and low muscle mass class 2, the prevalence of metabolic syndrome components were significantly increased, such as abdominal obesity, hypertension, hyperglycemia, and Low HDL cholesterolemia ( $P<0.05$ ), whereas in women, when the low muscle mass class was elevated, such as abdominal obesity, hypertension, hyperglycemia, and Low

Table 1. General characteristics of normal body mass index men

|  | Normal | Low muscle mass I | Low muscle mass II | $P$ value | $P$ value ${ }^{\text {a }}$ | $P$ value ${ }^{\text {b }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Age (yr) | $71.8 \pm 0.3$ | $73.3 \pm 0.5$ | $75.0 \pm 1.1$ | 0.001 | 0.001 | 0.008 |
| Ht (cm) | $164.6 \pm 0.3$ | $164.1 \pm 0.6$ | $162.5 \pm 0.8$ | 0.049 | 0.144 | 0.018 |
| Wt (kg) | $55.8 \pm 0.4$ | $57.7 \pm 0.6$ | $56.7 \pm 0.8$ | 0.016 | 0.004 | 0.379 |
| BMI (kg/m²) | $20.6 \pm 0.1$ | $21.4 \pm 0.1$ | $21.6 \pm 0.2$ | <0.001 | $<0.001$ | $<0.001$ |
| WC (cm) | $77.3 \pm 0.4$ | $81.2 \pm 0.7$ | $83.5 \pm 1.0$ | <0.001 | <0.001 | <0.001 |
| ASM (kg) | $18.6 \pm 0.1$ | $17.0 \pm 0.2$ | $15.2 \pm 0.3$ | <0.001 | <0.001 | <0.001 |
| ASM/Wt (\%) | $33.5 \pm 0.1$ | $29.5 \pm 0.1$ | $26.7 \pm 0.2$ | <0.001 | <0.001 | <0.001 |
| SBP (mmHg) | $128.0 \pm 1.1$ | $131.0 \pm 2.2$ | $138.8 \pm 3.6$ | 0.008 | 0.024 | 0.007 |
| DBP (mmHg) | $75.8 \pm 0.6$ | $76.7 \pm 1.0$ | $80.1 \pm 2.1$ | 0.144 | 0.161 | 0.062 |
| FBS (mg/dL) | $99.4 \pm 1.2$ | $111.1 \pm 2.9$ | $104.3 \pm 3.3$ | <0.001 | $<0.001$ | 0.398 |
| TG (mg/dL) | $121.7 \pm 5.1$ | $145.1 \pm 8.1$ | $148.1 \pm 14.8$ | 0.023 | 0.006 | 0.168 |
| HDLC (mg/dL) | $52.6 \pm 0.7$ | $51.0 \pm 1.6$ | $48.5 \pm 1.6$ | 0.051 | 0.128 | 0.028 |
| Alcohol None Moderate Heavy | $\begin{aligned} & 29.8(2.4) \\ & 25.2(2.3) \\ & 45.0(2.7) \end{aligned}$ | $\begin{aligned} & 36.5(4.8) \\ & 23.9(4.3) \\ & 39.6(5.1) \end{aligned}$ | $\begin{aligned} & 42.7(10.0) \\ & 43.2(10.7) \\ & 14.0(6.1) \end{aligned}$ | 0.024 | 0.102 | 0.009 |
| Smoking None Past Current | $\begin{aligned} & 13.0(1.8) \\ & 24.1(2.6) \\ & 62.9(2.6) \end{aligned}$ | $\begin{aligned} & 15.2(3.5) \\ & 28.8(4.4) \\ & 56.0(4.9) \end{aligned}$ | $\begin{aligned} & 16.4(5.9) \\ & 26.4(8.2) \\ & 57.2(9.7) \end{aligned}$ | 0.751 | 0.425 | 0.854 |
| Exercise <br> None Irregular Regular | $\begin{aligned} & 77.1(2.3) \\ & 12.1(1.9) \\ & 10.9(1.7) \end{aligned}$ | $\begin{array}{r} 81.1(4.3) \\ 9.7(3.1) \\ 9.2(3.4) \end{array}$ | $\begin{array}{r} 85.6(6.3) \\ 3.2(3.2) \\ 11.2(5.8) \end{array}$ | 0.647 | 0.455 | 0.424 |

Calculated by complex sample general linear model and complex sample logistic regression analysis. Values are expressed as mean $\pm$ standard error, estimated \% (standard error).
${ }^{\text {al }} P$-value for normal vs. low muscle mass I and II.
${ }^{\text {b }} P$-value for normal and low muscle mass grade I vs. low muscle mass grade II. Ht, height; Wt, weight; BMI, body mass index; WC, waist circumference; ASM, appendicular skeletal muscle mass; SBP, systolic blood pressure; DBP, diastolic blood pressure; FBS, fasting plasma sugar; TG, triglyceride; HDLC, high-density lipoprotein cholesterol.

HDL cholesterolemia were presented with statistically significant increases ( $P<0.05$ ). However, Low HDL cholesterolemia did not present any significant differences in both men and women ( $P=0.358, P=0.238$ ). In men, the proportion of metabolic syndrome was presented from $14.3 \%$ of the normal group, $23.3 \%$ of the group with low muscle mass class 1 and $46.3 \%$ of the group with low muscle mass class 2. Such results indicated that the metabolic syndrome proportion was high in the group with reduction of skeletal muscle mass ( $P<0.001$ ). Likewise in women, the prevalence of metabolic syndrome by each group was $36.8 \%, 59.0 \%$ and $64.2 \%$ respectively, presenting that the metabolic syndrome proportion was significantly increased as the low muscle mass class was progressed to higher level, similar to the men ( $P=0.002$ ) (Table 3).
When didn't adjust relevant variables (Model 1), the odds ratio for metabolic syndrome in men and women with low
muscle mass class 2 was presented as 5.15 and 3.08 ( $P<0.001$, $P=0.002$ ). When adjusted for age (Model 2 ), the likelihood of occurrence of metabolic syndrome in men was presented as 1.92 times greater and 5.83 times greater in those with low muscle mass class 1 and class 2 , compared to the normal group ( $P<0.001$ ), while in women, they were presented as 2.49 times greater and 3.12 times greater than the normal group ( $P=0.002$ ).
When adjusted for age and BMI (Model 3), the likelihood of occurrence of metabolic syndrome in men was presented as 1.53 times greater and 4.40 times greater in those with low muscle mass class 1 and 2 , presenting that it was statistically significantly increased ( $P=0.004$ ). In women, there were similar trends but with no statistical significance ( $P=0.074$ ). ( $P<0.001, P=0.002$ ). When additionally adjusted for smoking status, alcohol consumption, and exercise frequency (Model 4), it also presented a similar outcome to

Table 2. General characteristics of normal body mass index women

|  | Normal | Low muscle mass I | Low muscle mass II | Pvalue | Pvalue ${ }^{\text {a }}$ | $P$ value ${ }^{\text {b }}$ |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: |
| Age (yr) | $73.9 \pm 0.3$ | $73.4 \pm 0.8$ | $73.1 \pm 1.4$ | 0.763 | 0.497 | 0.630 |
| $\mathrm{Ht}(\mathrm{cm})$ | $150.3 \pm 0.3$ | $149.4 \pm 0.7$ | $149.0 \pm 1.2$ | 0.319 | 0.166 | 0.340 |
| Wt (kg) | $46.8 \pm 0.3$ | $48.8 \pm 0.5$ | $49.0 \pm 0.5$ | <0.001 | <0.001 | <0.001 |
| BMI (kg/m²) | $20.7 \pm 0.1$ | $21.8 \pm 0.1$ | $22.1 \pm 0.2$ | <0.001 | <0.001 | <0.001 |
| WC (cm) | $74.5 \pm 0.3$ | $79.2 \pm 1.0$ | $82.0 \pm 1.1$ | <0.001 | <0.001 | <0.001 |
| ASM (kg) | $12.7 \pm 0.1$ | $11.1 \pm 0.1$ | $1.0 \pm 0.1$ | <0.001 | $<0.001$ | <0.001 |
| ASM/wt (\%) | $27.3 \pm 0.1$ | $22.8 \pm 0.1$ | $20.3 \pm 0.1$ | <0.001 | $<0.001$ | $<0.001$ |
| SBP (mmHg) | $130.8 \pm 1.0$ | $134.0 \pm 1.8$ | $128.4 \pm 2.0$ | 0.092 | 0.195 | 0.207 |
| DBP (mmHg) | $74.7 \pm 0.5$ | $76.0 \pm 1.1$ | $74.1 \pm 2.4$ | 0.527 | 0.385 | 0.760 |
| FBS (mg/dL) | $98.8 \pm 1.0$ | $113.3 \pm 7.4$ | $114.1 \pm 14.8$ | 0.084 | 0.027 | 0.358 |
| TG (mg/dL) | $137.1 \pm 4.8$ | $154.6 \pm 10.2$ | $177.7 \pm 17.4$ | 0.036 | 0.043 | 0.032 |
| HDLC (mg/dL) | $51.5 \pm 0.6$ | $48.7 \pm 1.9$ | $49.2 \pm 5.8$ | 0.334 | 0.153 | 0.734 |
| Alcohol <br> None <br> Moderate Heavy | $\begin{array}{r} 65.8(2.6) \\ 27.0(2.5) \\ 7.2(1.2) \end{array}$ | $\begin{array}{r} 70.3(6.4) \\ 23.5(6.1) \\ 6.2(2.9) \end{array}$ | $\begin{array}{r} 62.3(9.5) \\ 37.7(9.5) \\ 0.0(0.0) \end{array}$ | 0.755 | 0.764 | 0.329 |
| Smoking None Past Current | $\begin{array}{r} 84.6(1.9) \\ 1.6(0.5) \\ 13.8(1.8) \end{array}$ | $\begin{array}{r} 85.4(6.2) \\ 1.5(1.5) \\ 13.1(6.1) \end{array}$ | $\begin{gathered} 84.9(10.3) \\ 0.0(0.0) \\ 15.1(10.3) \end{gathered}$ | 0.994 | 0.974 | 0.993 |
| Exercise None Irregular Regular | $\begin{aligned} & 74.9(2.2) \\ & 11.1(1.5) \\ & 14.0(1.9) \end{aligned}$ | $\begin{array}{r} 82.9(4.7) \\ 5.2(2.3) \\ 11.9(4.1) \\ \hline \end{array}$ | $\begin{array}{r} 94.0(5.8) \\ 0.0(0.0) \\ 6.0(5.8) \end{array}$ | 0.383 | 0.097 | 0.607 |

Calculated by complex sample general linear model and complex sample logistic regression analysis.
Values are expressed as mean $\pm$ standard error, estimated \% (standard error).
${ }^{\text {a }} P$ value for normal vs. low muscle mass I and II.
${ }^{\text {b }}$ Pvalue for normal and low muscle mass grade I vs. Iow muscle mass grade II. Ht, height; Wt, weight; BMI, body mass index; WC, waist circumference; ASM, appendicular skeletal muscle mass; SBP, systolic blood pressure; DBP, diastolic blood pressure; FBS, fasting plasma sugar; TG, triglyceride; HDLC, high-density lipoprotein cholesterol.

Table 3. Frequency of metabolic components according to muscle mass status

|  | Men |  |  |  | Women |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Normal | Low muscle mass I | Low muscle mass II | Pfor trend | Normal | Low muscle mass I | Low muscle mass II | Pfor trend |
| Abdominal obesity | 0.8 (0.5) | 5.8 (2.5) | 12.3 (5.4) | $<0.001$ | 22.8 (2.0) | 39.2 (7.3) | 67.4 (15.9) | $<0.001$ |
| Hypertension | 52.2 (2.7) | 57.7 (5.2) | 83.6 (6.8) | 0.004 | 62.1 (2.5) | 76.2 (5.5) | 61.6 (11.9) | 0.180 |
| Hyperglycemia | 31.8 (2.4) | 49.8 (5.1) | 62.8 (8.1) | $<0.001$ | 33.0 (2.4) | 48.9 (7.3) | 41.5 (11.4) | 0.029 |
| Hypertriglyceridemia | 24.6 (2.6) | 39.1 (4.9) | 47.1 (10.4) | 0.001 | 33.4 (2.3) | 58.3 (6.9) | 51.1 (10.5) | 0.001 |
| Low HDL Cholesterolemia | 21.4 (2.2) | 20.1 (3.7) | 33.0 (7.6) | 0.358 | 54.1 (2.4) | 61.5 (6.7) | 65.0 (15.1) | 0.238 |
| Metabolic syndrome | 14.3 (2.0) | 23.3 (4.0) | 46.3 (10.4) | <0.001 | 36.8 (2.3) | 59.0 (6.9) | 64.2 (16.4) | 0.002 |

Calculated by complex sample logistic regression analysis.
Values are expressed as estimated \% (standard error).
HDL, high-density lipoprotein.
those adjusted for age and $\mathrm{BMI}(P=0.002, P=0.123)$ (Table 4).

## DISCUSSION

According to this study which had analyzed the data of
the KNHANES, $18.8 \%$ of adult men $\geq 65$ years old with normal BMI was identified as low muscle mass class 1 and 5.2\% as low muscle mass class 2 . On the contrary, the low muscle mass class 1 and class 2 in women had shown $11.2 \%$ and $2.2 \%$ of prevalence respectively. When analyzed the

Table 4. Odds ratios of metabolic syndrome by muscle mass

|  | Men |  |  |  | Women |  |  |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | Normal | Low muscle mass I | Low muscle mass II | Pfor trend | Normal | Low muscle mass I | Low muscle mass II | Pfor trend |
| Model 1 | 1 | 1.81 | 5.15 | <0.001 | 1 | 2.48 | 3.08 | 0.002 |
| Model 2 | 1 | 1.92 | 5.83 | <0.001 | 1 | 2.49 | 3.12 | 0.002 |
| Model 3 | 1 | 1.53 | 4.40 | 0.004 | 1 | 1.68 | 1.91 | 0.074 |
| Model 4 | 1 | 1.63 | 5.53 | 0.002 | 1 | 1.73 | 1.55 | 0.123 |

Calculated by complex sample logistic regression analysis.
Values are presented as prevalence odds ratio.
Model 1: unadjusted, Model 2: adjusted for age, Model 3: adjusted for age and body mass index, Model 4: adjusted for age, body mass index, smoking status, alcohol consumption and exercise status.
aged subjects $\geq 65$ years old whose BMI was $\geq 23.0 \mathrm{~kg} / \mathrm{m}^{2}$ as collected from the data of the KNHANES conducted in the same period to this study, the men with low muscle mass class 1 was $39.5 \%$ and class 2 was $19.5 \%$ out of 883 male subjects. In comparison, out of 1,450 female subjects, $40.6 \%$ was low muscle mass class 1 and $14.5 \%$ was class 2 , suggesting the aged with normal BMI presented lower prevalence of low muscle mass than the aged with obesity.
In this study, when adjusted for age, the low skeletal muscle mass was an independent risk factor of metabolic syndrome in both men and women, and as the low muscle mass class was progressed, the likelihood of metabolic syndrome was eventually increased. These findings were similar results to the studies that had analyzed the relationship between sarcopenia and metabolic syndrome without limiting BMI presented up to now. Kim et al.[7] had defined and analyzed sarcopenia in their study, using the ASM adjusted for Wt in the selected subjects aged $\geq 20$ years old, also adjusted for BP, and presented that when adjusted for such as total cholesterol, fasting blood glucose and smoking status, the likelihood of developing metabolic syndrome in men with level 1 sarcopenia was increased by 3.5 times and in men with level 2 , it was increase by 4 times with statistical significance. In women, the odd ratio was 1.5 and 2.7 respectively. Such results were presented in similar pattern in a study conducted in the aged, and in a study that had identified the association between sarcopenia and cardiovascular diseases in the aged $\geq 65$ years old, sarcopenia had an independent association with cardiovascular diseases even if adjusted for the cardiovascular diseases-related variables.[10]
The pathophysiology of sarcopenia is not clear, yet, it has been known that it was developed by chronic inflammation status caused by reduction in producing testosterone,
cortisol, and vitamin D and increase of resistance to them, decrease in muscular cells, muscle protein degradation and abnormalities of cytokine secretion regulation.[11] In sarcopenia state, the muscle mass is reduced which is primarily involved in glucose metabolism therefore the insulin resistance is to be increased regardless obesity status, and consequently, the person becomes vulnerable to diabetes and various metabolic diseases. Furthermore, due to the reduction of muscle mass, basal metabolic rate and physical activities are reduced resulting in increase of body fat, in addition to increase prevalence of fall, functional disability, and metabolism disorder as well as cardiovascular diseases, which will lead to increase of mortality.[5]
According to the results of this study, it presented that the likelihood of metabolic syndrome prevalence is increased as the low muscle mass class is progressed more particularly in men. It is also similar to the previous study, so the relevance with hormones should be considered.[15] Testosterone in men increases muscle mass and strength to activate the catabolism and assimilation path of protein. On the other hand, estrogen of women is primarily known to be involved in adipocytes.[16] As the secretion of male hormones is decreased and the low muscle mass class is progressed, there is a potential for increased prevalence of metabolic syndrome.

As for the limits of this study, first, it is a cross-sectional study using the data of the KNHANES, and by this study alone, it would be difficult to find out the prevalence of low skeletal muscle mass, and it was not possible to identify the causal relationship between the low skeletal muscle mass and metabolic syndrome. Secondly, in the European Working Group on Sarcopenia in Older People (EWCSOP) recently presented, it suggests deterioration of muscular strength or reduction of activities of daily living in addition
to reduction in skeletal muscle mass as the conditions for defining sarcopenia.[17] Nonetheless, in this study, only the reduction of skeletal muscle mass was evaluated. But considering the correlation status of muscle mass and strength has been already well known,[18] if the low skeletal muscle mass was assessed by the criteria in accordance with this study, it is possible to predict that the muscle strength would also has been reduced. Thirdly, the skeletal muscle mass used in assessing sarcopenia is defined by a number of ways. The value obtained from dividing ASM by the square of $\mathrm{Ht}\left(\mathrm{ASM} / \mathrm{Ht} 2, \mathrm{~kg} / \mathrm{m}^{2}\right.$ ) is also one of the marker to assess sarcopenia. However, with the ASM adjusted for Ht , it is difficult to adjust for the Wt appropriately so that it is difficult to adequately compensate the increase of muscle mass happening along with the increase of body fat mass to the greater level.[19] Also, there was an analysis reported that the ASM adjusted for Ht recently is inadequate to assess the functional impairment in the aged.[20] Also there was a study reported that the indicator of ASM adjusted for Wt had deeper association with metabolic diseases compared to the value adjusted for Ht [9] Therefore, this study had evaluated skeletal muscle mass by using the indicator of ASM adjusted for Wt, which was expected to reflect metabolic diseases relatively well among a variety of defining methods. Fourth, the inflammatory proteins such as C-reactive protein (CPR) and various cytokines had been known to cause sarcopenia.[21] Also, chronic inflammatory conditions are appeared to have association with metabolic disorders.[22] In addition, there was a lack of analysis on the argument that change in sex hormones has effects on the development of metabolic syndrome. In future research, it would require assessment of muscle strength and activities of daily living, coping with limitations of this study, and so would be a study that considers association of various inflammatory substances.
This study has its significance in terms that it had shown the reduction of skeletal muscle mass was a major risk factors of metabolic syndrome associated with future cardiovascular events even in the aged with normal BMI. When the BMI is normal, there are many cases that the assessment of obesity is not fulfilled not only by the person but also in primary care, thus, it is easy to have delays of appropriate education and changes in the lifestyle.
However, like the results of this study, the risk of metabolic syndrome can be elevated depending on the muscle
mass, even if the BMI is normal. This indicates that there should be efforts to assess the muscle mass and obesity even in the aged with normal BMI, while since the skeletal muscle mass can be changed by adequate dietary and exercise methods, there should be educations available for those facts.

## REFERENCES

1. WHO Expert Consultation. Appropriate body-mass index for Asian populations and its implications for policy and intervention strategies. Lancet 2004;363:157-63.
2. Tanaka S, Togashi K, Rankinen T, et al. Is adiposity at normal body weight relevant for cardiovascular disease risk? Int J Obes Relat Metab Disord 2002;26:176-83.
3. Kwon HJ, Ha YC, Park HM. The Reference Value of Skeletal Muscle Mass Index for Defining the Sarcopenia of Women in Korea. J Bone Metab 2015;22:71-5.
4. 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 2011;12:249-56.
5. Lee JE, Kim KW, Paik NJ, et al. Evaluation of factors influencing grip strength in elderly koreans. J Bone Metab 2012; 19:103-10.
6. Metter EJ, Talbot LA, Schrager M, et al. Skeletal muscle strength as a predictor of all-cause mortality in healthy men. J Gerontol A Biol Sci Med Sci 2002;57:B359-65.
7. Kim KM, Lim S, Choi SH, et al. Cardiometabolic implication of sarcopenia: The Korea National Health and Nutrition Examination Study (KNHANES) 2008-2010. IJC Metab Endocr 2014;4:63-9.
8. Koo HS, Kim MJ, Kim KM, et al. Decreased muscle mass is not an independent risk factor for metabolic syndrome in Korean population aged 70 or older. Clin Endocrinol (Oxf) 2015;82:509-16.
9. Lim S, Kim JH, Yoon JW, et al. Sarcopenic obesity: prevalence and association with metabolic syndrome in the Korean Longitudinal Study on Health and Aging (KLoSHA). Diabetes Care 2010;33:1652-4.
10. Chin SO, Rhee SY, Chon S, et al. Sarcopenia is independently associated with cardiovascular disease in older Korean adults: the Korea National Health and Nutrition Examination Survey (KNHANES) from 2009. PLoS One 2013;8:e60119.
11. Evans WJ. Skeletal muscle loss: cachexia, sarcopenia, and inactivity. Am J Clin Nutr 2010;91:1123S-7S.
12. Janssen I, Heymsfield SB, Ross R. Low relative skeletal muscle mass (sarcopenia) in older persons is associated with functional impairment and physical disability. J Am Geriatr Soc 2002;50:889-96.
13. Grundy SM, Cleeman JI, Daniels SR, et al. Diagnosis and management of the metabolic syndrome: an American Heart Association/National Heart, Lung, and Blood Institute scientific statement: Executive Summary. Crit Pathw Cardiol 2005;4:198-203.
14. Lee SY, Park HS, Kim DJ, et al. Appropriate waist circumference cutoff points for central obesity in Korean adults. Diabetes Res Clin Pract 2007;75:72-80.
15. Di Monaco M, Castiglioni C, Vallero F, et al. Sarcopenia is more prevalent in men than in women after hip fracture: a cross-sectional study of 591 inpatients. Arch Gerontol Geriatr 2012;55:e48-52.
16. Blackman MR, Sorkin JD, Münzer T, et al. Growth hormone and sex steroid administration in healthy aged women and
men: a randomized controlled trial. JAMA 2002;288:228292.
17. Cruz-Jentoft AJ, Baeyens JP, Bauer JM, et al. Sarcopenia: European consensus on definition and diagnosis: Report of the European Working Group on Sarcopenia in Older People. Age Ageing 2010;39:412-23.
18. Reed RL, Pearlmutter L, Yochum K, et al. The relationship between muscle mass and muscle strength in the elderly. J Am Geriatr Soc 1991;39:555-61.
19. Newman AB, Kupelian V, Visser M, et al. Sarcopenia: alternative definitions and associations with lower extremity function. J Am Geriatr Soc 2003;51:1602-9.
20. Visser M. Towards a definition of sarcopenia-results from epidemiologic studies. J Nutr Health Aging 2009;13:713-6.
21. Abbatecola AM, Paolisso G, Fattoretti P, et al. Discovering pathways of sarcopenia in older adults: a role for insulin resistance on mitochondria dysfunction. J Nutr Health Aging 2011;15:890-5.
22. Jensen GL. Inflammation: roles in aging and sarcopenia. JPEN J Parenter Enteral Nutr 2008;32:656-9.
