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Measures of muscle mass and fat mass in the identification of  
metabolic abnormalities in older Korean adults



The Graduate School  
Yonsei University  
Department of Public Health

Measures of muscle mass and fat mass in the identification of  
metabolic abnormalities in older Korean adults

A Master's Thesis

Submitted to the Department of Public Health  
and the Graduate School of Yonsei University  
in partial fulfillment of the  
requirements for the degree of  
Master of Public Health

Ji Hye Park

June 2015

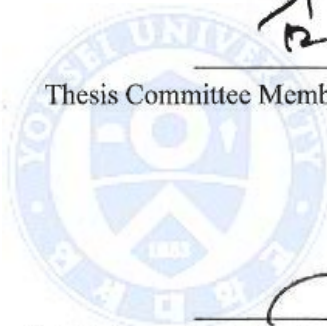
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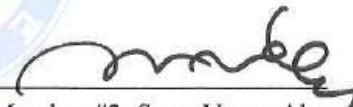


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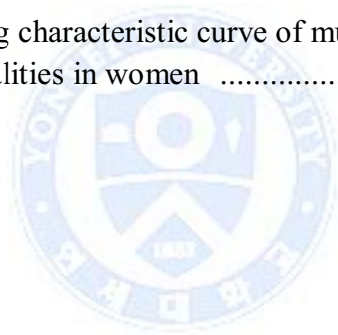
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## **ABSTRACT**

# **Measures of muscle mass and fat mass in the identification of metabolic abnormalities in older Korean adults**

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*The graduate School of Yonsei University*

(Directed by Professor Hyeon Chang Kim)

## **OBJECTIVES:**

We investigated the association of the sex-associated changes of muscle mass and fat mass with metabolic abnormalities in an older Korean population.

## **METHODS:**

We conducted a cross-sectional analysis of the baseline data from the cohort study conducted in the Korean Urban Rural Elderly (KURE) study, which is a population-based longitudinal study of health determinants among elderly persons aged 65 years or older (381 men, 747 women). Metabolic syndrome was defined according to the National Cholesterol Education Program's ATP-III criteria ( $\geq 3$  of the following abnormalities): waist circumference greater than 90 cm in men and 80 cm in women; serum triglycerides level of at least 150 mg/dL; high-density lipoprotein (HDL) cholesterol level of less than 40 mg/dL in men and 50 mg/dL in women; blood pressure of at least 130/85 mmHg; or serum glucose level of at least 100 mg/dL. The association between muscle and fat mass and metabolic syndrome was assessed by serial logistic regression models.

## **RESULTS:**

Fat mass was significantly associated with all components of the metabolic syndrome in both sexes. After adjustment for potential confounders including fat mass, muscle mass was associated with high blood pressure (ASM/Ht<sup>2</sup>; OR= 2.46, 95% CI = 1.61-3.75), low HDL cholesterol (ASM; OR= 1.91, 95% CI = 1.17-2.88 and ASM/Ht<sup>2</sup>; OR= 2.25, 95% CI = 1.49-3.38), high glucose (ASM; OR= 1.61, 95% CI = 1.05-2.48) and metabolic syndrome (ASM/Ht<sup>2</sup>; OR= 1.65, 95% CI = 1.12-

2.42) for women and low HDL cholesterol ( $ASM/Ht^2$ ; OR= 1.88, 95% CI = 1.01-3.49) for men.

## **CONCLUSIONS:**

In older persons, fat mass was associated with all of the metabolic syndrome components. In contrast, muscle mass was associated with all of the metabolic syndrome components in women, but not in men. More studies are needed to explain the sex difference of the associations.



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**Keywords:** Muscle mass, fat mass, metabolic abnormality, elderly

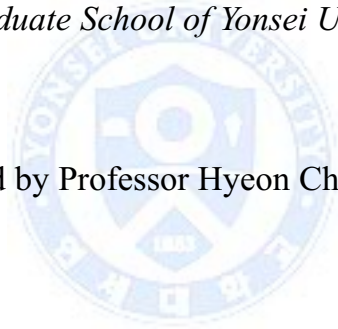
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## **I. INTRODUCTION**

### ***1. Background***

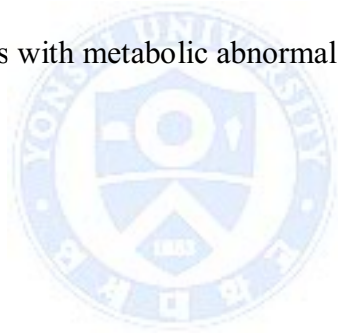
Metabolic syndrome is defined as a cluster of hypertension, hyperglycemia, dyslipidemia, and abdominal obesity ("Third Report of the National Cholesterol

Education Program (NCEP) Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults (Adult Treatment Panel III) final report" 2002). Metabolic syndrome is associated with cardiovascular disease which is the leading cause of mortality and morbidity . In Korea, the prevalence of metabolic syndrome, according to the National Cholesterol Education Program (NCEP)–Adult Treatment Panel (ATP) III, was 25.7 % in men and 31.9 % in women (Yoon et al. 2007) and the prevalence of metabolic syndrome were steadily increasing in elderly people (Ford, Giles and Dietz 2002; Park et al. 2007).

In addition, previous studies have suggested that the effects of metabolic syndrome may depend on age (Roriz-Cruz et al. 2007). Insulin resistance, one of the components of metabolic syndrome, has been considered as a contributing factor to age-related muscle mass loss, which is causally related to decline in functional ability. Moreover, older individuals tend to have a greater proportion of fat than younger people with the same BMI. Both cross-sectional and longitudinal studies have shown that age-related body composition changes, such as fat mass increase and muscle mass decrease (Baumgartner et al. 1995; Forbes 1999). The abdominal obesity including fat mass was well known to be strongly associated with metabolic syndrome (Bosy-Westphal et al. 2006b) and lower muscle mass, termed sarcopenia, was also associated with metabolic syndrome (Ishii et al. 2014).

## **2. Objective**

There is no unanimous view about the standard criteria of sarcopenia to apply to define low muscle mass, since classification of sarcopenia differs by ethnic groups and equipment for measuring the muscle mass (Alexandre Tda et al. 2014). Therefore, we assessed the association between absolute muscle mass and metabolic syndrome components without classifying sarcopenia among older Korean adults. We also investigated the association of the sex-associated changes of muscle mass and fat mass with metabolic abnormalities.

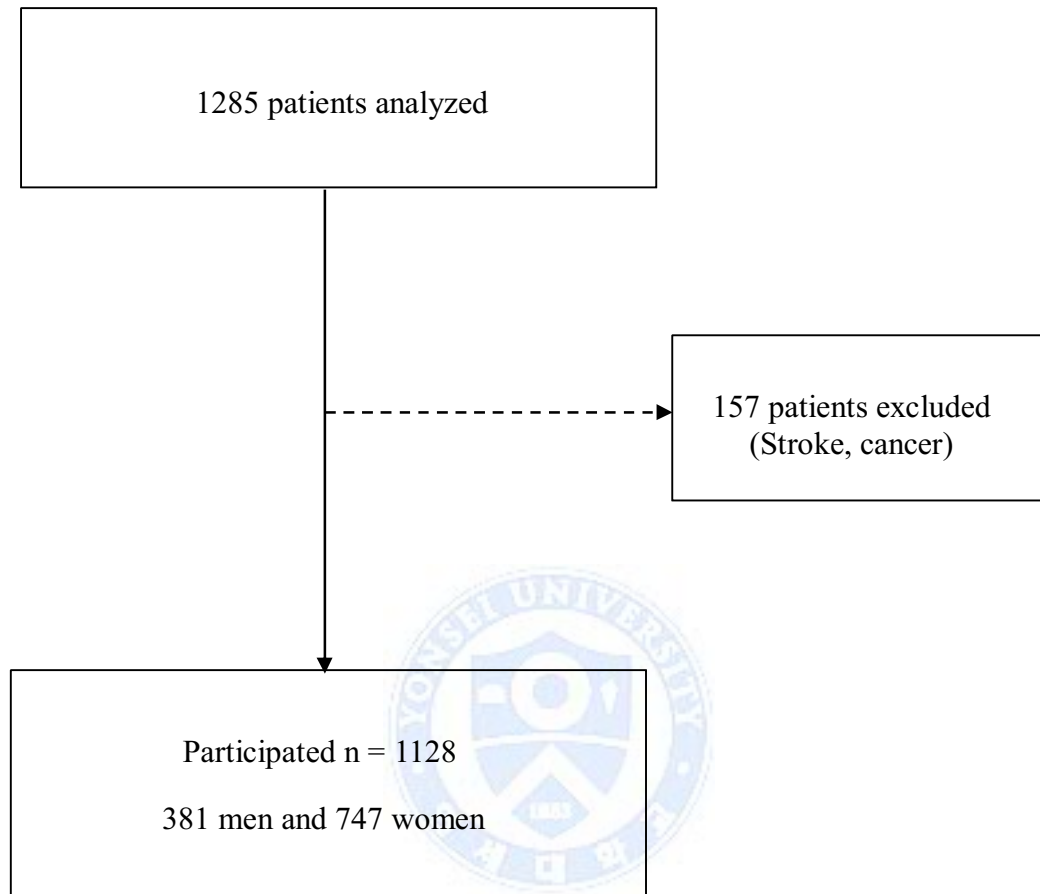


## II. METHODS

### *1. Study population*

The study is conducted using baseline data collected from Korean elderly participating in the Korean Urban Rural Elderly (KURE) study (Lee et al. 2014). The KURE study is a community-based prospective cohort study on health, aging, and common geriatric disorders of Korean elderly persons aged at least 65 years. To construct a cohort reflecting both urban and rural areas, we selected two representative communities in the country.

Between July and December 2014, a bioelectrical impedance ancillary study was performed for 1285 permanent residents. After excluding 175 participants with past history of cancer or stroke, 1128 participants were eligible for the current cross-sectional analysis (Figure 1). Among them, 760 participants were measured for anthropometric parameters and examined for fasting blood test in 2014, and 368 participants were measured for anthropometric parameters in 2014 and examined for fasting blood test in 2012. All participants provided written informed consents, and the study protocol was approved by the Institutional Review Board of Severance Hospital, Yonsei University College of Medicine.



**Figure1. Flowchart of the selection criteria for the final study population**



## **2. *Measurements***

### **1) Questionnaire**

Participants were individually interviewed using standardized questionnaires to obtain information about their general characteristics, medical history, medication use, and lifestyle behaviors. Trained interviewers carried out the questionnaire surveys according to the predefined protocol, and double-checked whether responses were inappropriate or missing. Smoking status was classified as current smokers or nonsmokers (past smokers or those who had never smoked). Alcohol consumption was categorized as regular alcohol drinking or other (participants who drink less than once a week or not at all). Physical activity was categorized as regular exercise or no exercise.

### **2) Physical Examination**

We measured height and weight with subjects in light clothing and calculated body mass index (BMI) as weight in kilograms divided by the square of height in meters ( $\text{kg/m}^2$ ). Waist circumference was measured between the lower borders of the rib cage and the iliac crest with a measuring tape (SECA-201; SECA, Hamburg, Germany). Resting blood pressure was measured twice by an automatic sphygmomanometer (Dinamap 1846 SX/P; GE Healthcare, Waukesha, WI, USA)

with the participant in the sitting position at least 5 minute intervals. If the difference between the first and second measurement was more than 10 mmHg for either systolic or diastolic blood pressure, a third measurement was performed, and the last two measurements were averaged for analyses. Muscle mass and fat mass were measured by bioelectrical impedance analysis (BIA) using an Inbody 720 machine (Biospace, Seoul, Korea). Appendicular skeletal muscle mass (ASM) was derived as the sum of the muscle mass of the four limbs (Cruz-Jentoft et al. 2010). We used ASM divided by weight (ASM/Wt) and by height squared (ASM/Ht<sup>2</sup>) as muscle mass indices; fat mass divided by weight (Body fat/Wt) and by height squared (Body fat/Ht<sup>2</sup>) as fat mass indices. The results of our study did not differ significantly when divided by weight and by height squared. Therefore, only ASM/Ht<sup>2</sup> and Body fat/Ht<sup>2</sup> indices were used in the analysis. Grip strength was measured with a hand dynamometer with participants seated, their elbow by their side and flexed to right angles, and a neutral wrist position. The measurements were conducted in each hand with 20 seconds rest intervals, and the mean value of four measures was used in the analysis.

### 3) Laboratory Assays

Blood samples were collected from the antecubital vein after at least an 8 hour fast. Enzymatic methods were applied to measure total cholesterol, HDL

cholesterol, and triglycerides and fasting blood glucose level were measured by colorimetry method with Auto Analyzer (ADVIA 1800; Siemens Healthcare Diagnostics Inc., Deerfield, IL, USA). Low-density lipoprotein cholesterol was calculated using the Friedewald' method (Friedewald, Levy and Fredrickson 1972).

### ***3. Definition of metabolic abnormalities***

Metabolic syndrome was defined based on the National Cholesterol Education Program Adult Treatment Panel III (NCEP-ATP III) criteria (Alberti et al. 2009). The presence of any three of the following five abnormalities constitutes a diagnosis of metabolic syndrome: (i) waist circumference >90 cm in men and >80 cm in women; (ii) elevated triglycerides with fasting plasma triglycerides  $\geq 150$  mg/dL; (iii) low HDL cholesterol with fasting HDL cholesterol <40 mg/dL in men and <50 mg/dL in women; (iv) elevated blood pressure with systolic blood pressure  $\geq 130$  mmHg and/or diastolic blood pressure  $\geq 85$  mmHg; (v) elevated fasting plasma glucose with fasting plasma glucose  $\geq 100$  mg/dL.

#### 4. *Statistical Analysis*

Differences in subject characteristics between men and women were examined using Student's *t*-test, ANOVA or Wilcoxon rank-sum test (for continuous variables) and chi-square test (for categorical variables). The correlation between muscle mass and fat mass and other variables were evaluated by the Pearson's correlation coefficient controlling for age, smoking status, physical activity, and alcohol intake. Also, the Spearman's correlation coefficients were used for skewed variables. We employed logistic regression analysis to evaluate the association between muscles mass and fat mass. Multiple logistic regression analysis was used to assess the odds ratio for the individual metabolic abnormalities per one unit increase in the muscle mass and fat mass. We applied the following serial models: age-adjusted (model 1); age, potential confounders such as smoking, drinking and physical activity-adjusted (model 2). In the final model, age, potential confounders and fat mass or muscle mass (ASM with corresponding body fat and ASM/Ht<sup>2</sup> with corresponding body fat/Ht<sup>2</sup>) were included (model 3). Furthermore, the receiver operating characteristic (ROC) analysis was used to compare the discriminative power of muscle mass and fat mass. All statistical analyses were performed using SAS version 9.2 (SAS Institute, Cary, NC, USA), and statistical significance was defined as a two-sided *p*-value less than  $\leq 0.05$ .

### **III. RESULTS**

#### ***1. Characteristics of study populations***

General characteristics for men and women participants are shown in Table 1. The variables were significantly different between men and women, with the exception of triglycerides and insulin. Men had higher muscle mass, blood pressure and fasting glucose, tended to smoke more, and drank more alcohol than women. Women had higher fat mass, total cholesterol, HDL cholesterol and LDL cholesterol, and more physical activity than men/higher physical activity level than men.

**Table 1. Characteristics of study populations**

Variables	Men (n=381)	Women (n=747)	<i>p</i> -value
Age, year	72.5 ± 4.2	71.1 ± 4.4	<.001
Height, cm	164.9 ± 5.7	152.7 ± 5.3	<.001
Weight, kg	65.0 ± 8.4	57.1 ± 7.9	<.001
Body mass index, kg/m <sup>2</sup>	23.9 ± 2.7	24.5 ± 3.1	0.001
Waist circumference, cm	86.0 ± 8.7	82.6 ± 8.9	<.001
ASM, kg	20.5 ± 2.7	14.5 ± 2.0	<.001
Body fat, kg	16.4 ± 5.3	20.1 ± 5.8	<.001
ASM/Ht	7.5 ± 0.7	6.2 ± 0.6	<.001
Body fat/Ht	6.0 ± 2.0	8.6 ± 2.5	<.001
Grip strength, kg	32.0 ± 5.8	19.9 ± 4.4	<.001
Lifestyle characteristics			
Current smoker, %	265 (69.7)	12 (1.6)	<.001
Drinker, %	268 (70.3)	180 (24.1)	<.001
Physical inactivity, %	129 (33.9)	296 (39.6)	<.001
Metabolic risk factors			
Systolic blood pressure, mmHg	128.9 ± 13.6	127.0 ± 15.8	0.040
Diastolic blood pressure, mmHg	74.7 ± 8.5	73.4 ± 8.6	0.012
Total cholesterol, mg/dL	172.5 ± 33.1	184.9 ± 34.4	<.001
HDL cholesterol, mg/dL	47.4 ± 11.6	51.1 ± 12.2	<.001
LDL cholesterol, mg/dL	100.3 ± 28.9	108.7 ± 29.8	<.001
Triglycerides, mg/dL	109 [83-150]	112 [83-153]	0.767
Fasting glucose, mg/dL	98 [91-108]	94 [88-104]	0.027
Insulin, uIU/L	5.1 [3.3-8.1]	6.1 [4.1-9.5]	0.124

Data are expressed as means ± standard deviation, median [inter quartile range] and number (%)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein;

LDL, low density lipoprotein

## ***2. Description of study populations by tertiles of muscle mass and body fat***

Description of men and women by tertile of ASM and body fat are shown in Table 2 and 3. Men and women in the lowest tertile of ASM were older and shorter, had a lower weight, BMI, waist circumference and body fat, and had a lower grip strength compared with those in the highest tertile (Table 2). Among the highest tertile of ASM, low HDL cholesterol, high glucose and metabolic syndrome were significantly more prevalent in women ( $p = 0.02$ ,  $p = 0.01$ , and  $p < .001$ , respectively), while metabolic syndrome was significantly more prevalent in men ( $p = 0.004$ ). Men and women in the highest tertile of body fat had a higher weight, BMI and waist circumference compared with those in the lowest tertile. Among the highest tertile of body fat, metabolic syndrome and its components were significantly more prevalent in both sexes (Table 3).

**Table 2. Description of men and women by tertiles of muscle mass**

Variables	Men				Women			
	Tertile 1	Tertile 2	Tertile 3	<i>p</i> -value	Tertile 1	Tertile 2	Tertile 3	<i>p</i> -value
Age, year	74.3 ± 4.0	72.1 ± 4.1	71.1 ± 3.8	<.001	72.6 ± 4.8	71.1 ± 4.1	69.6 ± 3.7	<.001
Height, cm	160.3 ± 4.5	164.8 ± 3.7	169.5 ± 4.7	<.001	148.4 ± 4.0	152.7 ± 4.0	156.9 ± 4.1	<.001
Weight, kg	58.3 ± 6.8	64.3 ± 4.9	72.3 ± 6.6	<.001	50.9 ± 5.6	56.7 ± 5.4	63.6 ± 6.9	<.001
Body mass index, kg/m <sup>2</sup>	22.7 ± 2.8	23.7 ± 2.4	25.2 ± 2.4	<.001	23.2 ± 2.9	24.4 ± 2.9	25.9 ± 3.0	<.001
Waist circumference, cm	82.1 ± 8.7	85.4 ± 7.9	90.4 ± 7.4	<.001	79.1 ± 8.4	82.1 ± 8.4	86.7 ± 8.2	<.001
ASM, kg	17.8 ± 1.5	20.4 ± 0.6	23.4 ± 1.8	<.001	17.7 ± 5.2	20.0 ± 5.3	22.6 ± 5.9	<.001
ASM/Ht	6.9 ± 0.6	7.5 ± 0.4	8.2 ± 0.6	<.001	8.1 ± 2.5	8.6 ± 2.5	9.2 ± 2.5	<.001
Body fat, kg	15.0 ± 5.7	15.9 ± 4.4	18.2 ± 5.2	<.001	12.3 ± 1.0	14.4 ± 0.5	16.7 ± 1.1	<.001
Body fat/Ht	5.9 ± 2.2	5.9 ± 1.7	6.4 ± 1.9	0.082	5.6 ± 0.4	6.2 ± 0.3	6.8 ± 0.4	<.001
Grip strength, kg	29.5 ± 4.6	31.8 ± 5.6	34.7 ± 5.9	<.001	18.1 ± 3.9	19.3 ± 4.0	22.2 ± 4.3	<.001
Metabolic abnormality								
High blood pressure	66 (53.2)	75 (58.6)	78 (61.4)	0.411	123 (49.6)	129 (51.6)	146 (59.4)	0.072
Low HDL cholesterol	28 (22.4)	44 (34.4)	48 (37.8)	0.022	116 (46.8)	142 (56.8)	144 (58.5)	0.018
High triglycerides	24 (19.2)	33 (25.8)	39 (30.7)	0.108	58 (23.4)	65 (26.0)	69 (28.1)	0.494
High glucose	50 (40.0)	56 (43.8)	60 (47.2)	0.511	67 (27.2)	88 (35.2)	98 (39.8)	0.010
Metabolic syndrome	33 (26.4)	47 (36.7)	59 (46.5)	0.004	88 (35.5)	111 (44.4)	136 (55.3)	<.001

Data are expressed as means ± standard deviation, median [inter quartile range] and number (%)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein



**Table 3. Description of men and women by tertiles of body fat**

Variables	Men				Women			
	Tertile 1	Tertile 2	Tertile 3	<i>p</i> -value	Tertile 1	Tertile 2	Tertile 3	<i>p</i> -value
Age, year	72.7 ± 4.4	72.4 ± 3.8	72.4 ± 4.3	0.742	71.0 ± 4.2	71.3 ± 4.9	71.0 ± 4.0	0.549
Height, cm	164.9 ± 5.5	164.4 ± 6.0	165.4 ± 5.7	0.365	152.1 ± 5.3	152.6 ± 5.1	153.3 ± 5.6	0.062
Weight, kg	57.6 ± 6.1	64.9 ± 5.0	72.2 ± 6.5	<.001	49.6 ± 4.5	56.7 ± 4.2	64.7 ± 6.0	<.001
Body mass index, kg/ m <sup>2</sup>	21.2 ± 2.0	24.0 ± 1.4	26.4 ± 1.7	<.001	21.4 ± 1.8	24.4 ± 1.5	27.6 ± 2.2	<.001
Waist circumference, cm	78.1 ± 7.7	86.2 ± 4.8	93.3 ± 5.3	<.001	74.6 ± 6.3	83.1 ± 5.2	90.0 ± 7.1	<.001
Body fat, kg	10.7 ± 2.2	16.0 ± 1.3	22.2 ± 3.0	<.001	14.0 ± 2.7	19.8 ± 1.4	26.4 ± 3.7	<.001
Body fat/Ht	3.9 ± 0.9	5.9 ± 0.6	8.1 ± 1.2	<.001	6.1 ± 1.3	8.5 ± 0.8	11.3 ± 1.7	<.001
ASM, kg	19.7 ± 2.6	20.5 ± 2.3	21.3 ± 2.9	<.001	13.6 ± 1.7	14.4 ± 1.8	15.4 ± 2.0	<.001
ASM/Ht	7.2 ± 0.8	7.6 ± 0.5	7.8 ± 0.7	<.001	5.9 ± 0.5	6.2 ± 0.5	6.5 ± 0.6	<.001
Grip strength, kg	31.6 ± 5.7	32.2 ± 5.1	32.2 ± 6.5	0.659	20.0 ± 4.0	19.7 ± 4.7	19.9 ± 4.4	0.808
Metabolic abnormality								
High blood pressure	50 (40.0)	68 (55.3)	101 (77.1)	<.001	94 (38.2)	128 (52.0)	176 (69.9)	<.001
Low HDL cholesterol	23 (18.4)	43 (34.7)	54 (41.2)	<.001	109 (44.3)	147 (59.8)	146 (57.9)	0.001
High triglycerides	17 (13.6)	37 (29.8)	42 (32.1)	<.001	47 (19.1)	73 (39.7)	72 (28.6)	0.013
High glucose	38 (30.4)	49 (39.5)	79 (60.3)	<.001	61 (24.8)	88 (35.8)	104 (41.3)	<.001
Metabolic syndrome	13 (10.4)	37 (29.8)	89 (67.9)	<.001	52 (21.1)	121 (49.2)	162 (64.3)	<.001

Data are expressed as means ± standard deviation, median [inter quartile range] and number (%)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

### **3. Correlations between muscle mass and fat mass with metabolic variables**

The associations between muscle mass and fat mass with metabolic variables after adjusting for age, smoking, drinking and physical activity are shown in Table 4 and 5. For man, ASM was significantly positively correlated with body fat ( $r = 0.237, p < .001$ ), insulin ( $r = 0.147, p = 0.004$ ) and grip strength ( $r = 0.271, p < .001$ ), was significantly negatively correlated with total cholesterol ( $r = -0.112, p = 0.030$ ) and HDL cholesterol ( $r = -0.187, p < .001$ );  $ASM/Ht^2$  was significantly positively correlated with body fat ( $r = 0.269, p < .001$ ), body fat/ $Ht^2$  ( $r = 0.216, p < .001$ ), blood pressure ( $r = 0.104, p = 0.045$ ), insulin ( $r = 0.220, p < .001$ ) and grip strength ( $r = 0.199, p < .001$ ), was significantly negatively correlated with HDL cholesterol ( $r = -0.210, p < .001$ ). Body fat and body fat/ $Ht^2$  was significantly positively correlated with  $ASM/Ht^2$ , HDL cholesterol, blood pressure, triglycerides, fasting glucose and insulin, was significantly negatively correlated with HDL cholesterol (Table 4). For women, association muscle mass and fat mass was stronger than in men ( $r = 0.237, p < .001$  for men and  $r = 0.417, p < .001$  for women). Both muscle mass and fat mass indices were significantly positively correlated with blood pressure (ASM excepted), triglycerides, fasting glucose and insulin, were significantly negatively correlated with HDL cholesterol (Table 5).

**Table 4. Correlation analysis between muscle mass and fat mass with metabolic variables in men**

Variables	ASM		Body fat		ASM/Ht		Body fat/Ht	
	r	p-value	r	p-value	r	p-value	r	p-value
Height, cm	<b>0.696</b>	<b>&lt;.001</b>	0.060	0.246	<b>0.196</b>	<b>&lt;.001</b>	<b>-0.153</b>	<b>0.003</b>
Weight, kg	<b>0.759</b>	<b>&lt;.001</b>	<b>0.785</b>	<b>&lt;.001</b>	<b>0.697</b>	<b>&lt;.001</b>	<b>0.677</b>	<b>&lt;.001</b>
Body mass index, kg/m <sup>2</sup>	<b>0.414</b>	<b>&lt;.001</b>	<b>0.832</b>	<b>&lt;.001</b>	<b>0.653</b>	<b>&lt;.001</b>	<b>0.846</b>	<b>&lt;.001</b>
Waist circumference, cm	<b>0.459</b>	<b>&lt;.001</b>	<b>0.785</b>	<b>&lt;.001</b>	<b>0.507</b>	<b>&lt;.001</b>	<b>0.738</b>	<b>&lt;.001</b>
ASM, kg	-	-	<b>0.237</b>	<b>&lt;.001</b>	<b>0.837</b>	<b>&lt;.001</b>	0.079	0.127
Body fat, kg	<b>0.237</b>	<b>&lt;.001</b>	-	-	<b>0.269</b>	<b>&lt;.001</b>	<b>0.975</b>	<b>&lt;.001</b>
ASM/Ht	<b>0.837</b>	<b>&lt;.001</b>	<b>0.269</b>	<b>&lt;.001</b>	-	-	<b>0.216</b>	<b>&lt;.001</b>
Body fat/Ht	0.079	0.127	<b>0.975</b>	<b>&lt;.001</b>	<b>0.216</b>	<b>&lt;.001</b>	-	-
Grip strength, kg	<b>0.271</b>	<b>&lt;.001</b>	-0.001	0.988	<b>0.199</b>	<b>&lt;.001</b>	-0.047	0.363
Blood pressure, mmHg	0.053	0.308	<b>0.150</b>	<b>0.004</b>	<b>0.104</b>	<b>0.045</b>	<b>0.159</b>	<b>0.002</b>
Total cholesterol, mg/dL	<b>-0.112</b>	<b>0.030</b>	-0.033	0.529	-0.085	0.102	-0.016	0.753
HDL cholesterol, mg/dL	<b>-0.187</b>	<b>&lt;.001</b>	<b>-0.321</b>	<b>&lt;.001</b>	<b>-0.210</b>	<b>&lt;.001</b>	<b>-0.311</b>	<b>&lt;.001</b>
Triglycerides, mg/dL*	0.068	0.186	<b>0.337</b>	<b>&lt;.001</b>	0.079	0.126	<b>0.325</b>	<b>&lt;.001</b>
Fasting glucose, mg/dL*	0.021	0.690	<b>0.240</b>	<b>&lt;.001</b>	0.035	0.498	<b>0.236</b>	<b>&lt;.001</b>
Insulin, uIU/L*	<b>0.147</b>	<b>0.004</b>	<b>0.644</b>	<b>&lt;.001</b>	<b>0.220</b>	<b>&lt;.001</b>	<b>0.631</b>	<b>&lt;.001</b>

Adjustment for age, smoking status, physical activity, and alcohol intake

Abbreviations: ASM, Appendicular skeletal muscle mass; HDL, high density lipoprotein

Correlation coefficients (r) and p-values were calculated with Pearson's (for normally distributed variables) or

\*Spearman's (for non-normally distributed variables) correlation coefficients.

**Table 5. Correlation between muscle mass and fat mass with metabolic variables in women**

Variables	ASM		Body fat		ASM/Ht		Body fat/Ht	
	r	p-value	r	p-value	r	p-value	r	p-value
Height, cm	<b>0.717</b>	<b>&lt;.001</b>	<b>0.081</b>	<b>0.027</b>	<b>0.279</b>	<b>&lt;.001</b>	<b>-0.154</b>	<b>&lt;.001</b>
Weight, kg	<b>0.739</b>	<b>&lt;.001</b>	<b>0.889</b>	<b>&lt;.001</b>	<b>0.747</b>	<b>&lt;.001</b>	<b>0.789</b>	<b>&lt;.001</b>
Body mass index, kg/m <sup>2</sup>	<b>0.411</b>	<b>&lt;.001</b>	<b>0.908</b>	<b>&lt;.001</b>	<b>0.651</b>	<b>&lt;.001</b>	<b>0.930</b>	<b>&lt;.001</b>
Waist circumference, cm	<b>0.433</b>	<b>&lt;.001</b>	<b>0.798</b>	<b>&lt;.001</b>	<b>0.538</b>	<b>&lt;.001</b>	<b>0.773</b>	<b>&lt;.001</b>
ASM, kg	-	-	<b>0.417</b>	<b>&lt;.001</b>	<b>0.867</b>	<b>&lt;.001</b>	<b>0.242</b>	<b>&lt;.001</b>
Body fat, kg	<b>0.417</b>	<b>&lt;.001</b>	-	-	<b>0.512</b>	<b>&lt;.001</b>	<b>0.969</b>	<b>&lt;.001</b>
ASM/Ht	<b>0.867</b>	<b>&lt;.001</b>	<b>0.512</b>	<b>&lt;.001</b>	-	-	<b>0.439</b>	<b>&lt;.001</b>
Body fat/Ht	<b>0.242</b>	<b>&lt;.001</b>	<b>0.969</b>	<b>&lt;.001</b>	<b>0.439</b>	<b>&lt;.001</b>	-	-
Grip strength, kg	<b>0.359</b>	<b>&lt;.001</b>	-0.002	0.964	<b>0.273</b>	<b>&lt;.001</b>	<b>-0.077</b>	<b>0.037</b>
Blood pressure, mmHg	0.035	0.347	<b>0.112</b>	<b>0.002</b>	<b>0.102</b>	<b>0.005</b>	<b>0.126</b>	<b>0.001</b>
Total cholesterol, mg/dL	-0.070	0.057	-0.044	0.232	<b>-0.079</b>	<b>0.032</b>	-0.040	0.273
HDL cholesterol, mg/dL	<b>-0.218</b>	<b>&lt;.001</b>	<b>-0.175</b>	<b>&lt;.001</b>	<b>-0.250</b>	<b>&lt;.001</b>	<b>-0.156</b>	<b>&lt;.001</b>
Triglycerides, mg/dL*	<b>0.107</b>	<b>0.004</b>	<b>0.197</b>	<b>&lt;.001</b>	<b>0.140</b>	<b>&lt;.001</b>	<b>0.183</b>	<b>&lt;.001</b>
Fasting glucose, mg/dL*	<b>0.172</b>	<b>&lt;.001</b>	<b>0.169</b>	<b>&lt;.001</b>	<b>0.142</b>	<b>&lt;.001</b>	<b>0.135</b>	<b>&lt;.001</b>
Insulin, uIU/L*	<b>0.252</b>	<b>&lt;.001</b>	<b>0.440</b>	<b>&lt;.001</b>	<b>0.290</b>	<b>&lt;.001</b>	<b>0.413</b>	<b>&lt;.001</b>

Adjustment for age, smoking status, physical activity, and alcohol intake

Abbreviations: ASM, Appendicular skeletal muscle mass; HDL, high density lipoprotein

Correlation coefficients (r) and *p*-values were calculated with Pearson's (for normally distributed variables) or

\*Spearman's (for non-normally distributed variables) correlation coefficients.

#### **4. Correlations between muscle mass and fat mass**

The relationships between muscle mass and fat mass were also presented using scatter plots, separately for men and women (Figure 2). In men, ASM was significantly and positively correlation with body fat ( $r = 0.237, p < .001$ ), and  $ASM/Ht^2$  was significantly and positively correlation with body fat/Ht ( $r = 0.216, p < .001$ ). In women, ASM was significantly and positively correlation with body fat ( $r = 0.417, p < .001$ ), and  $ASM/Ht^2$  was significantly and positively correlation with body fat/Ht ( $r = 0.512, p < .001$ ). These results provided evidence that muscle mass is strongly correlated with fat mass in women than in men.



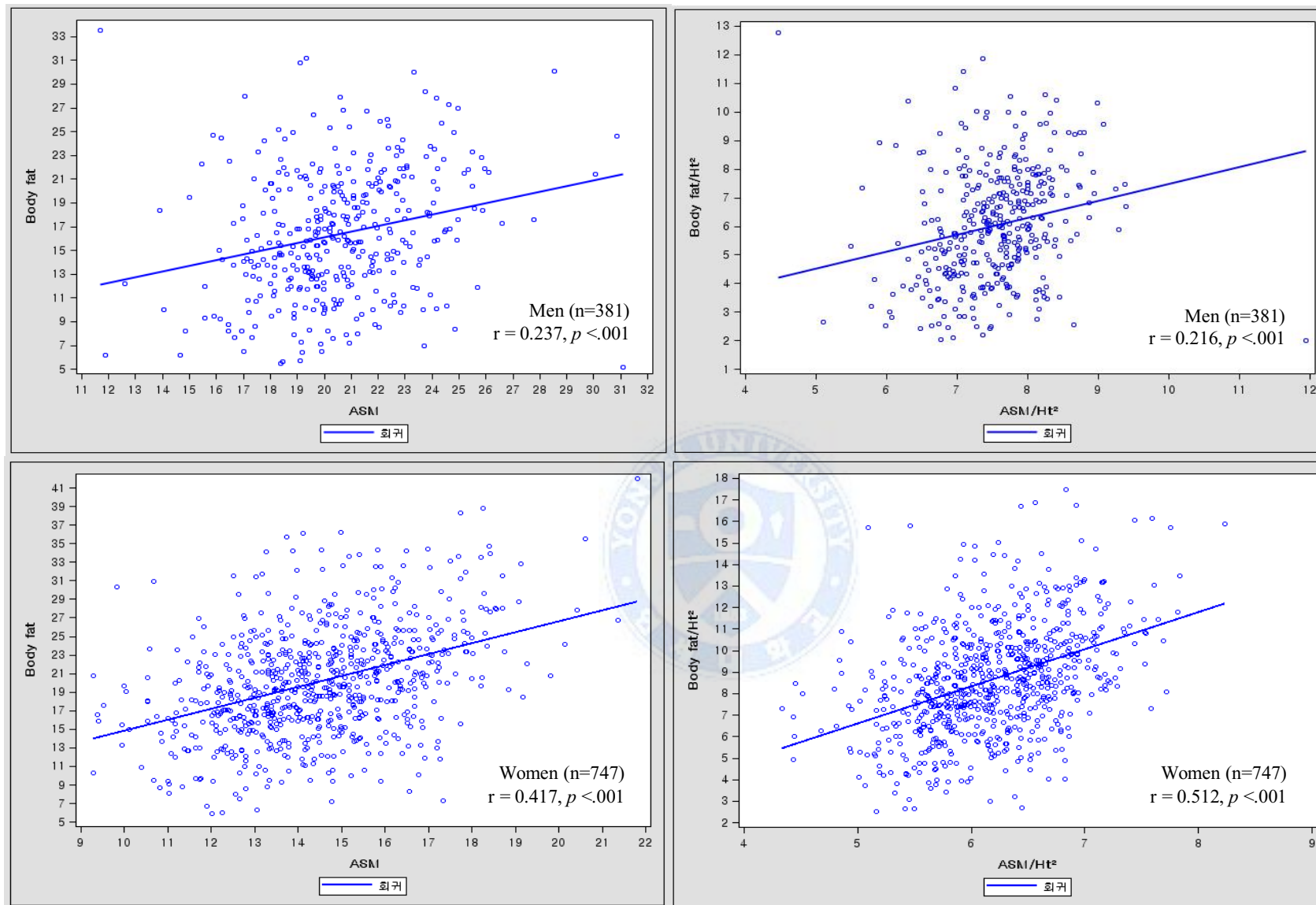


Figure.2 The relationship between muscle mass and fat mass in men and women

## **5. Association between muscle mass and fat mass and metabolic abnormalities**

Table 6, 7, 8 and 9 shows association between tertile of muscle mass and fat mass and metabolic abnormalities using multiple logistic regression analysis in each sex. In men, body fat and body fat/Ht were associated with all of the metabolic abnormalities, and further adjustment for muscle mass and potential confounders were significantly associated with all of the metabolic abnormalities. The highest tertile of body fat and body fat/Ht were 17.29 and 16.43 times, respectively, more likely to have an increased risk of metabolic syndrome than those in lowest tertile. In contrast, ASM and ASM/Ht<sup>2</sup> were associated with low HDL cholesterol and metabolic syndrome, but after adjustment for body fat and potential confounders the association was significant with low HDL cholesterol (ASM; OR = 2.24, 95% CI = 1.48-3.38) (Table 6 and 7). In women, body fat and body fat/Ht were associated with all of the metabolic abnormalities, but further adjustment for muscle mass and potential confounders was significantly associated with high blood pressure, high glucose and metabolic syndrome. The highest tertile of body fat and body fat/Ht were 5.14 and 4.15 times, respectively, more likely to have an increased risk of metabolic syndrome than those in lowest tertile. In contrast, ASM was associated with high blood pressure, low HDL cholesterol, high glucose and metabolic syndrome, and further adjustment for body fat mass and potential confounders was significantly associated with low HDL cholesterol (OR = 1.90, 95% CI 1.26-2.27), high glucose (OR = 1.59, 95% CI = 1.03-2.45) and metabolic syndrome (OR = 1.99, 95% CI = 1.28-3.09) (Table 8). ASM/Ht<sup>2</sup> was associated with all of the metabolic abnormalities, but further adjustment for body fat/Ht<sup>2</sup> and potential confounders was significantly associated with high blood pressure (OR = 2.39, 95% CI = 1.56-3.66), low HDL cholesterol (OR = 2.24, 95% CI = 1.48-3.38) and metabolic syndrome (OR = 2.47, 95%

CI = 1.60-3.81) (Table 9).





**Table 6. Logistic regression models of ASM and body fat mass for metabolic abnormality in men**

Men (n=381)	ASM		Body fat	
	Tertile 2	Tertile 3	Tertile 2	Tertile 3
	OR (95% CI)		OR (95% CI)	
High blood pressure				
Model 1	1.40 (0.83, 2.34)	1.66 (0.97, 2.83)	<b>1.90 (1.14, 3.15)</b>	<b>5.21 (3.01, 9.00)</b>
Model 2	1.42 (0.84, 2.39)	1.57 (0.91, 2.71)	<b>1.96 (1.17, 3.30)</b>	<b>5.43 (3.10, 9.48)</b>
Model 3	1.24 (0.72, 2.15)	1.02 (0.56, 1.86)	<b>1.96 (1.16, 3.30)</b>	<b>5.43 (3.06, 9.64)</b>
Low HDL cholesterol				
Model 1	<b>1.83 (1.04, 3.24)</b>	<b>2.13 (1.20, 3.81)</b>	<b>2.34 (1.31, 4.21)</b>	<b>3.10 (1.75, 5.48)</b>
Model 2	<b>1.85 (1.04, 3.31)</b>	<b>2.44 (1.33, 4.45)</b>	<b>2.42 (1.33, 4.40)</b>	<b>3.29 (1.83, 5.90)</b>
Model 3	1.72 (0.95, 3.09)	<b>1.88 (1.01, 3.49)</b>	<b>2.26 (1.23, 4.12)</b>	<b>2.81 (1.54, 5.12)</b>
High triglycerides				
Model 1	1.30 (0.71, 3.40)	1.58 (0.86, 2.92)	<b>2.68 (1.41, 5.11)</b>	<b>2.97 (1.58, 5.60)</b>
Model 2	1.27 (0.69, 2.35)	1.66 (0.89, 3.10)	<b>2.83 (1.47, 5.45)</b>	<b>3.12 (1.64, 5.93)</b>
Model 3	1.24 (0.67, 2.31)	1.35 (0.71, 2.56)	<b>2.81 (1.46, 5.41)</b>	<b>3.06 (1.59, 5.89)</b>
High glucose				
Model 1	1.14 (0.68, 1.90)	1.29 (0.76, 2.18)	1.49 (0.88, 2.51)	<b>3.46 (2.06, 5.81)</b>
Model 2	1.13 (0.67, 1.88)	1.27 (0.75, 2.17)	1.46 (0.86, 2.48)	<b>3.44 (3.04, 5.79)</b>
Model 3	1.04 (0.61, 1.76)	0.93 (0.53, 1.64)	1.46 (0.86, 2.49)	<b>3.44 (2.01, 5.88)</b>
Metabolic syndrome				
Model 1	1.69 (0.97, 2.92)	<b>2.57 (1.47, 4.50)</b>	<b>3.66 (1.83, 7.31)</b>	<b>18.24 (9.23, 36.07)</b>
Model 2	1.66 (0.96, 2.89)	<b>2.68 (1.52, 4.73)</b>	<b>3.75 (1.87, 7.54)</b>	<b>19.10 (9.57, 38.12)</b>
Model 3	1.59 (0.85, 2.98)	1.54 (0.80, 2.99)	<b>3.57 (1.77, 7.19)</b>	<b>17.29 (8.60, 34.75)</b>

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

The lowest tertile (tertile1) was used as reference group.

Model1: adjusted for age.

Model2: adjusted for age, smoking, drinking and physical activity.

Model3: adjusted for age, smoking, drinking, physical activity, and body fat (ASM).

**Table 7. Logistic regression models of ASM/Ht<sup>2</sup> and Body fat/Ht<sup>2</sup> for metabolic abnormality in men**

Men (n=381)	ASM/Ht		Body fat/Ht	
	Tertile 2	Tertile 3	Tertile 2	Tertile 3
	OR (95% CI)		OR (95% CI)	
High blood pressure				
Model 1	<b>1.65 (0.99, 2.76)</b>	<b>1.74 (1.03, 2.94)</b>	<b>1.64 (1.00, 2.71)</b>	<b>5.70 (3.25, 10.00)</b>
Model 2	1.58 (0.94, 2.68)	1.68 (0.98, 2.88)	<b>1.68 (1.00, 2.82)</b>	<b>6.06 (3.41, 10.78)</b>
Model 3	1.21 (0.69, 2.11)	1.02 (0.56, 1.85)	1.68 (0.99, 2.85)	<b>6.06 (3.34, 11.01)</b>
Low HDL cholesterol				
Model 1	1.45 (0.83, 2.54)	<b>1.79 (1.02, 3.13)</b>	<b>3.06 (1.70, 5.52)</b>	<b>3.22 (1.79, 5.81)</b>
Model 2	1.62 (0.91, 2.88)	<b>1.99 (1.11, 3.58)</b>	<b>3.29 (1.79, 6.05)</b>	<b>3.38 (1.85, 6.17)</b>
Model 3	1.41 (0.78, 2.53)	1.52 (0.83, 2.80)	<b>3.00 (1.62, 5.55)</b>	<b>2.85 (1.53, 5.31)</b>
High triglycerides				
Model 1	1.74 (0.95, 3.16)	1.41 (0.76, 2.62)	<b>2.61 (1.39, 4.89)</b>	<b>2.68 (1.43, 5.03)</b>
Model 2	<b>1.89 (1.02, 3.48)</b>	1.53 (0.81, 2.89)	<b>2.70 (1.42, 5.12)</b>	<b>2.81 (1.48, 5.33)</b>
Model 3	1.66 (0.89, 3.09)	1.20 (0.62, 2.31)	<b>2.67 (1.39, 5.11)</b>	<b>2.76 (1.43, 5.36)</b>
High glucose				
Model 1	<b>1.79 (1.07, 2.99)</b>	1.54 (0.91, 2.61)	<b>1.78 (1.06, 3.00)</b>	<b>3.33 (1.98, 5.61)</b>
Model 2	<b>1.86 (1.10, 3.13)</b>	1.57 (0.92, 2.68)	<b>1.71 (1.01, 2.90)</b>	<b>3.28 (1.94, 5.56)</b>
Model 3	1.57 (0.92, 2.70)	1.13 (0.64, 2.00)	<b>1.72 (1.01, 2.95)</b>	<b>3.32 (1.92, 5.75)</b>
Metabolic syndrome				
Model 1	<b>1.98 (1.15, 3.43)</b>	<b>2.71 (1.55, 4.73)</b>	<b>3.91 (1.97, 7.75)</b>	<b>18.12 (9.14, 35.92)</b>
Model 2	<b>2.15 (1.23, 3.75)</b>	<b>2.91 (1.65, 5.15)</b>	<b>3.89 (1.94, 7.78)</b>	<b>18.79 (9.39, 37.60)</b>
Model 3	1.64 (0.86, 3.10)	1.58 (0.82, 3.06)	<b>3.55 (1.76, 7.15)</b>	<b>16.43 (8.13, 33.18)</b>

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

The lowest tertile (tertile1) was used as reference group.

Model1: adjusted for age.

Model2: adjusted for age, smoking, drinking and physical activity.

Model3: adjusted for age, smoking, drinking, physical activity, and body fat/Ht (ASM/Ht ).

**Table 8. Logistic regression models of ASM and body fat mass for metabolic abnormality in women**

Women (n=747)	ASM		Body fat	
	Tertile 2	Tertile 3	Tertile 2	Tertile 3
	OR (95% CI)		OR (95% CI)	
High blood pressure				
Model 1	1.26 (0.88, 1.81)	<b>2.01 (1.37, 2.94)</b>	<b>1.77 (1.24, 2.56)</b>	<b>3.87 (2.65, 5.65)</b>
Model 2	1.33 (0.92, 1.92)	<b>2.21 (1.49, 3.26)</b>	<b>1.76 (1.22, 2.53)</b>	<b>3.85 (2.63, 5.62)</b>
Model 3	1.05 (0.71, 1.55)	1.36 (0.89, 2.07)	<b>1.55 (1.06, 2.25)</b>	<b>3.00 (2.00, 4.50)</b>
Low HDL cholesterol				
Model 1	<b>1.65 (1.15, 2.38)</b>	<b>1.95 (1.34, 2.84)</b>	<b>1.86 (1.30, 2.66)</b>	<b>1.34 (1.22, 2.48)</b>
Model 2	<b>1.66 (1.15, 2.40)</b>	<b>2.17 (1.47, 3.19)</b>	<b>1.84 (1.28, 2.64)</b>	<b>1.75 (1.22, 2.52)</b>
Model 3	<b>1.56 (1.07, 2.27)</b>	<b>1.90 (1.26, 2.87)</b>	<b>1.62 (1.12, 2.36)</b>	1.36 (0.92, 2.01)
High triglycerides				
Model 1	1.19 (0.79, 1.80)	1.39 (0.91, 2.12)	<b>1.82 (1.20, 2.76)</b>	<b>1.69 (1.11, 2.58)</b>
Model 2	1.17 (0.77, 1.77)	1.46 (0.95, 2.25)	<b>1.79 (1.18, 2.73)</b>	<b>1.69 (1.11, 1.59)</b>
Model 3	1.07 (0.70, 1.64)	1.23 (0.78, 1.94)	<b>1.69 (1.10, 2.59)</b>	1.59 (0.94, 2.35)
High glucose				
Model 1	<b>1.58 (1.07, 2.33)</b>	<b>2.07 (1.39, 3.09)</b>	<b>1.70 (1.15, 2.51)</b>	<b>2.14 (1.46, 3.13)</b>
Model 2	<b>1.57 (1.06, 2.32)</b>	<b>2.06 (1.37, 3.08)</b>	<b>1.74 (1.18, 2.58)</b>	<b>2.18 (1.48, 3.21)</b>
Model 3	1.39 (0.93, 2.08)	<b>1.59 (1.03, 2.45)</b>	<b>1.52 (1.02, 2.27)</b>	<b>1.65 (1.09, 2.50)</b>
Metabolic syndrome				
Model 1	<b>1.68 (1.16, 2.45)</b>	<b>3.07 (2.08, 4.53)</b>	<b>3.69 (2.48, 5.49)</b>	<b>6.89 (4.60, 10.32)</b>
Model 2	<b>1.79 (1.23, 2.63)</b>	<b>3.54 (2.36, 5.29)</b>	<b>3.77 (2.52, 5.64)</b>	<b>7.11 (4.72, 10.72)</b>
Model 3	1.37 (0.91, 2.05)	<b>1.99 (1.28, 3.09)</b>	<b>3.22 (2.14, 4.87)</b>	<b>5.14 (3.33, 7.92)</b>

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

The lowest tertile (tertile1) was used as reference group.

Model1: adjusted for age.

Model2: adjusted for age, smoking, drinking and physical activity.

Model3: adjusted for age, smoking, drinking, physical activity, and body fat (ASM).

**Table 9. Logistic regression models of ASM/Ht<sup>2</sup> and Body fat/Ht<sup>2</sup> for metabolic abnormality in women**

Women (n=747)	ASM/Ht		Body fat/Ht	
	Tertile 2	Tertile 3	Tertile 2	Tertile 3
	OR (95% CI)		OR (95% CI)	
High blood pressure				
Model 1	1.94 (1.34, 2.80)	3.24 (2.21, 4.76)	2.14 (1.49, 3.07)	3.53 (2.43, 5.14)
Model 2	2.04 (1.41, 2.97)	3.58 (2.41, 5.31)	2.13 (1.48, 3.07)	3.46 (2.38, 5.05)
Model 3	1.64 (1.11, 2.42)	2.39 (1.56, 3.66)	1.79 (1.23, 2.61)	2.48 (1.64, 3.73)
Low HDL cholesterol				
Model 1	1.58 (1.11, 2.27)	2.23 (1.54, 3.23)	2.41 (1.68, 3.46)	1.72 (1.20, 2.46)
Model 2	1.66 (1.15, 2.39)	2.44 (1.66, 3.56)	2.39 (1.66, 3.45)	1.74 (1.21, 2.50)
Model 3	1.58 (1.09, 2.30)	2.24 (1.48, 3.38)	2.04 (1.40, 2.97)	1.25 (0.84, 1.87)
High triglycerides				
Model 1	1.39 (0.92, 2.11)	1.60 (1.05, 2.43)	1.87 (1.23, 2.84)	1.83 (1.20, 2.78)
Model 2	1.42 (0.93, 2.16)	1.66 (1.09, 2.54)	1.84 (1.21, 2.81)	1.84 (1.20, 2.81)
Model 3	1.29 (0.84, 1.99)	1.41 (0.89, 2.23)	1.67 (1.08, 2.57)	1.51 (0.95, 2.41)
High glucose				
Model 1	1.43 (0.98, 2.09)	1.68 (1.14, 2.47)	1.89 (1.29, 2.77)	1.88 (1.28, 2.77)
Model 2	1.43 (0.97, 2.10)	1.67 (1.13, 2.47)	1.92 (1.30, 2.82)	1.93 (1.31, 2.85)
Model 3	1.28 (0.86, 1.90)	1.36 (0.89, 2.07)	1.73 (1.16, 2.57)	1.58 (1.04, 2.42)
Metabolic syndrome				
Model 1	1.90 (1.30, 2.76)	3.67 (2.50, 5.41)	4.38 (2.95, 6.52)	6.00 (4.02, 8.96)
Model 2	2.05 (1.40, 3.00)	4.19 (2.81, 6.25)	4.45 (2.98, 6.64)	6.07 (4.04, 9.11)
Model 3	1.52 (1.02, 2.28)	2.47 (1.60, 3.81)	3.71 (2.46, 5.60)	4.15 (2.69, 6.40)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

The lowest tertile (tertile1) was used as reference group.

Model1: adjusted for age.

Model2: adjusted for age, smoking, drinking and physical activity.

Model3: adjusted for age, smoking, drinking, physical activity, and body fat/Ht (ASM/Ht ).

**6. The areas under the curves of muscle mass and fat mass in the prediction of metabolic abnormalities**

The areas under the curves (AUC) of muscle mass and fat mass in the prediction of metabolic abnormalities are shown in Table 10, 11 and Figure 3, 4. In men, the AUC of fat mass was greater than that of muscle mass in the prediction of all of the metabolic abnormalities. The AUCs for body fat and body fat /Ht for identifying High blood pressure were 0.692 (95% CI = 0.638-0.746) and 0.689 (95% CI = 0.635-0.743); low HDL cholesterol were 0.640 (95% CI = 0.582-0.697) and 0.627 (95% CI = 0.569-0.684); high triglycerides were 0.616 (95% CI = 0.556-0.677) and 0.607 (95% CI = 0.547-0.667); high glucose were 0.650 (95% CI = 0.595-0.705) and 0.645 (95% CI = 0.589-0.701); metabolic syndrome were 0.813 (95% CI = 0.768-0.857) and 0.799 (95% CI = 0.754-0.845), respectively (Table 10 and Figure 3). In women, the AUC of fat mass was greater than that of muscle mass in the prediction of high blood pressure, high triglycerides and metabolic syndrome. The AUCs for body fat and body fat /Ht for identifying high blood pressure were 0.660 (95% CI = 0.621-0.699) and 0.659 (95% CI = 0.620-0.698); high triglycerides were 0.573 (95% CI = 0.527-0.618) and 0.570 (95% CI = 0.524-0.616); metabolic syndrome were 0.717 (95% CI = 0.680-0.753) and 0.700 (95% CI = 0.663-0.737), respectively. The AUC of fat mass and muscle mass were showed similar in the prediction of low HDL cholesterol and high glucose. The AUCs for ASM/Ht and body fat for identifying low HDL cholesterol were 0.581 (95% CI = 0.524-0.607) and 0.572 (95% CI = 0.530-0.614), respectively. The AUCs for ASM and body fat for identifying high glucose were 0.584 (95% CI = 0.541-0.672) and 0.591 (95% CI = 0.548-0.633), respectively (Table 11 and Figure 4).

**Table 10. Comparison of areas under ROC curve for different muscle mass and fat mass by metabolic abnormalities in men**

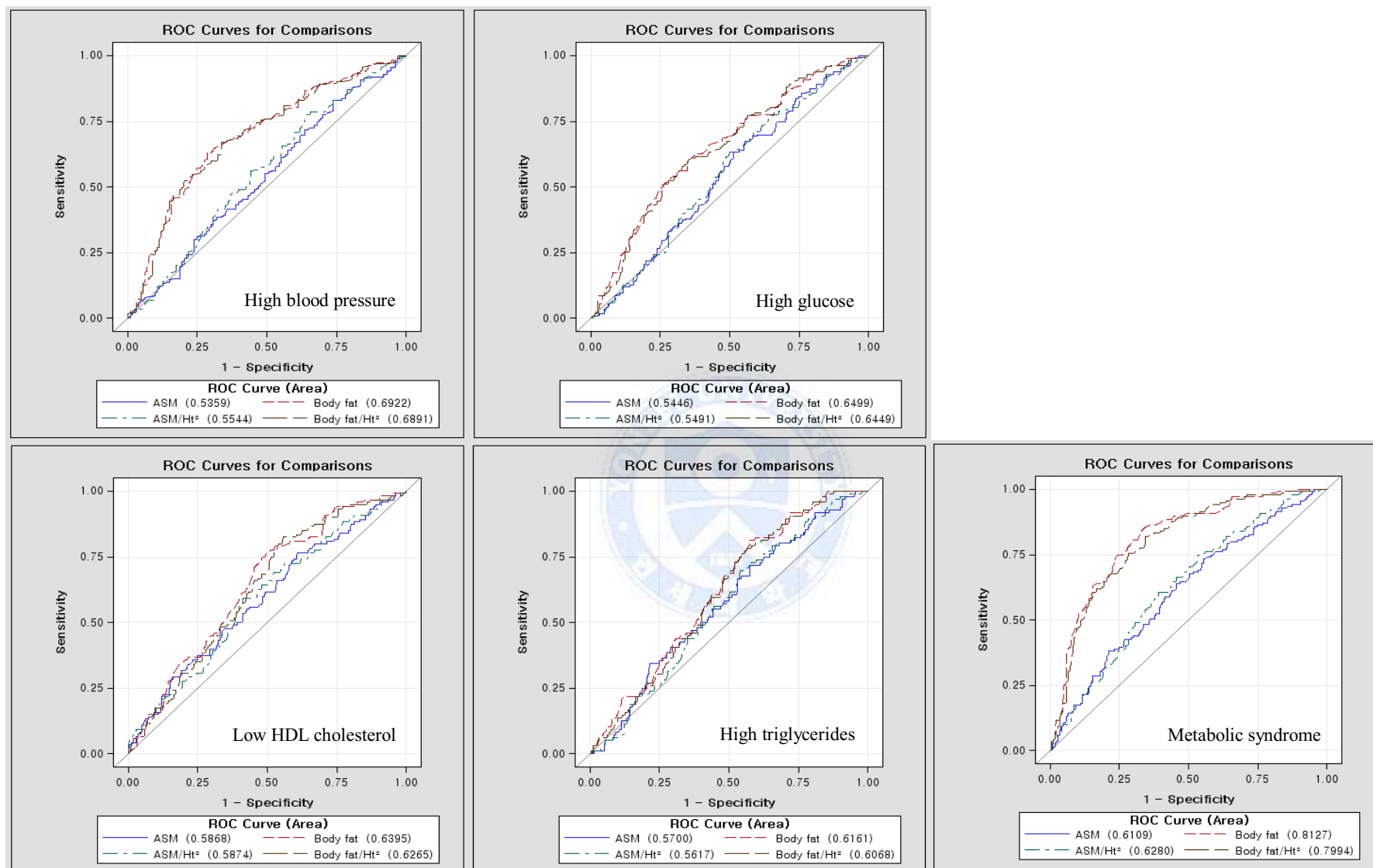
Men (n=381)	Areas under ROC curve for 95% CI				
	High blood pressure	Low HDL cholesterol	High triglycerides	High glucose	Metabolic syndrome
ASM	0.536(0.477-0.595)	0.587(0.525-0.649)	0.570(0.506-0.634)	0.545(0.487-0.603)	0.611(0.553-0.669)
Body fat	0.692(0.638-0.746)	0.640(0.582-0.697)	0.616(0.556-0.677)	0.650(0.595-0.705)	0.813(0.768-0.857)
ASM/Ht	0.554(0.495-0.614)	0.587(0.526-0.648)	0.562(0.499-0.625)	0.549(0.491-0.607)	0.628(0.571-0.685)
Body fat/Ht	0.689(0.635-0.743)	0.627(0.569-0.684)	0.607(0.547-0.667)	0.645(0.589-0.701)	0.799(0.754-0.845)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein

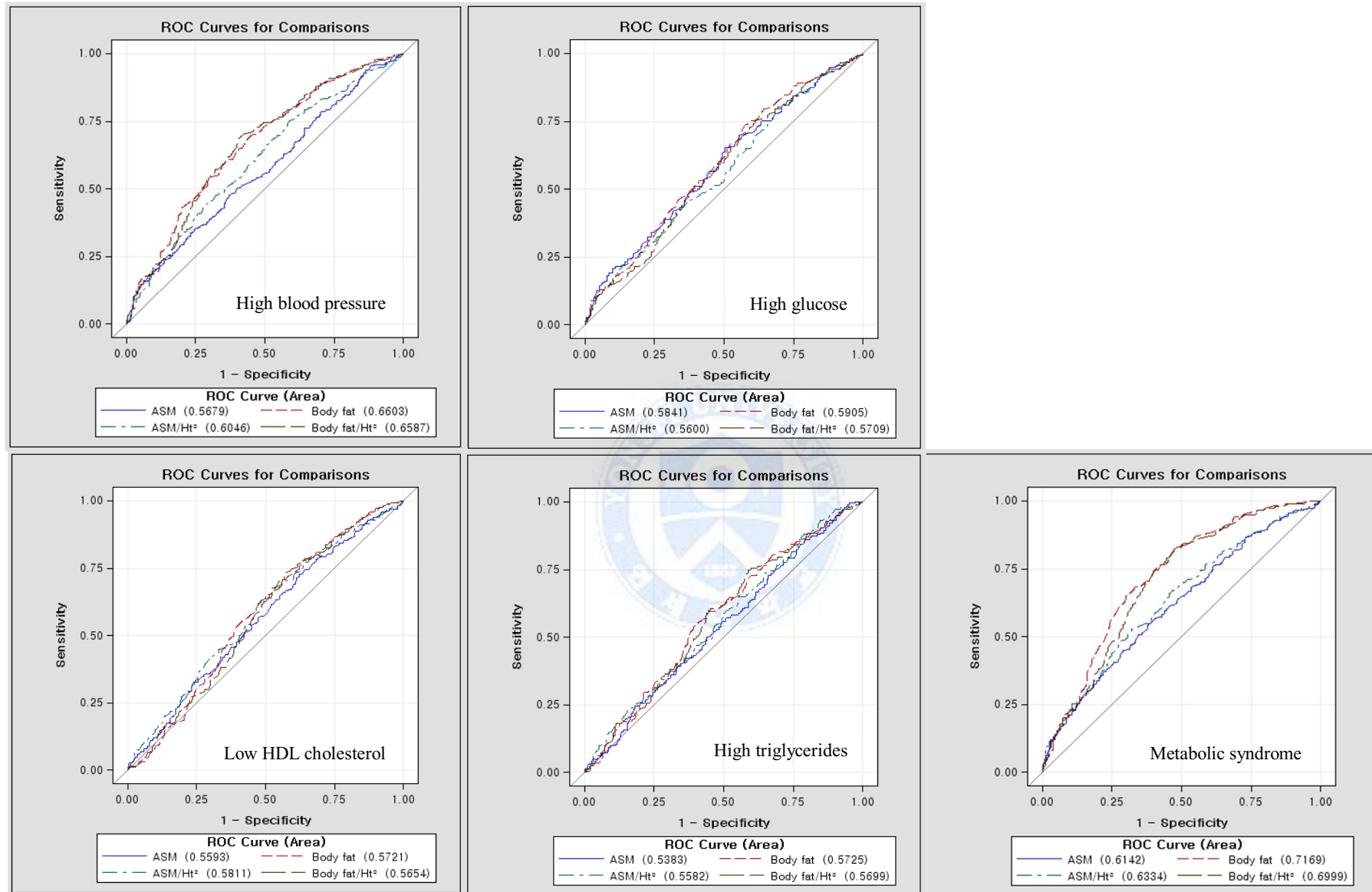
**Table 11. Comparison of areas under ROC curve for different muscle mass and fat mass by metabolic abnormalities in women**

Women (n=747)	Areas under ROC curve for 95% CI				
	High blood pressure	Low HDL cholesterol	High triglycerides	High glucose	Metabolic syndrome
ASM	0.568(0.527-0.609)	0.559(0.518-0.601)	0.538(0.492-0.585)	0.584(0.541-0.672)	0.614(0.574-0.655)
Body fat	0.660(0.621-0.699)	0.572(0.530-0.614)	0.573(0.527-0.618)	0.591(0.548-0.633)	0.717(0.680-0.753)
ASM/Ht	0.605(0.564-0.645)	0.581(0.540-0.622)	0.558(0.512-0.605)	0.560(0.517-0.603)	0.633(0.594-0.673)
Body fat/Ht	0.659(0.620-0.698)	0.565(0.524-0.607)	0.570(0.524-0.616)	0.571(0.528-0.614)	0.700(0.663-0.737)

Abbreviations: ASM, appendicular skeletal muscle; HDL, high density lipoprotein



**Figure.3 Receiver operating characteristic curve of muscle mass and fat mass and metabolic abnormalities in men**



**Figure.4 Receiver operating characteristic curve of muscle mass and fat mass and metabolic abnormalities in women**



## IV. DISCUSSION

### *1. Summary of finding*

The present study investigated that fat mass and muscle mass were associated with the metabolic syndrome along with its components in Korean older adults. We observed that higher fat mass was associated with increased risk of metabolic syndrome along with its components in both men and women. Furthermore, higher muscle mass was associated with increased risk of high blood pressure, low HDL cholesterol, high glucose and metabolic syndrome after adjustment for body fat and potential confounders only in women. Muscle mass is strongly correlated with fat mass in women than in men.

### *2. Comparison with previous studies*

In the elderly population, body composition such as fat mass and muscle mass, gradually changes with age even if the body weight remains unchanged (Gallagher et al. 2000; Kim et al. 2014). Previous studies have proven that fat mass is associated with inflammatory markers and metabolic abnormalities (Bosy-Westphal et al. 2006a; Forouhi, Sattar and McKeigue 2001). Consistent with those studies, our study showed that fat mass was related to metabolic abnormalities, independent of muscle mass and other potential confounders.

Meanwhile, previous studies have reported that low muscle mass reduces the intensity and endurance of physical activity (Wannamethee and Atkins 2015). These changes may increase of obesity and obesity-relates metabolic abnormalities in older people (Ishii et al. 2014; Karakelides and Nair 2005) and muscular strength was inversely associated with incident

metabolic syndrome (Jurca et al. 2005). Furthermore, both obesity and sarcopenia are associated with metabolic disorders and are important causes of disability, morbidity and mortality (Stephen and Janssen 2009; Wannamethee and Atkins 2015). However, our study showed that the positive associations between muscle mass and high blood pressure, low HDL cholesterol, high glucose and metabolic syndrome were observed only in women. These results show that women with high muscle mass have an especially greater risk of metabolic abnormalities than those with lower muscle mass, but this is not consistent with previous studies.

### ***3. Possible mechanism***

One of the possible underlying factors is validation of a BIA equation to predict muscle mass and fat mass. The BIA is simple, noninvasive, relatively inexpensive, easy-to-use method of estimating body composition. Numerous studies have developed equations for estimating lean body mass from BIA measurements (Bosaeus et al. 2014; Rangel Peniche, Raya Giorguli and Aleman-Mateo 2015). However, to ensure that reliable BIA measurements are obtained, several factors such as hydration status, food intake, and exercise must be controlled (Thibault, Genton and Pichard 2012).

Another possible underlying factor is the age-specific effects of metabolic syndrome. In middle aged populations, metabolic syndrome has been proven a relevant determinant of association with several outcomes, including cardiovascular and cerebrovascular morbidity and mortality (Thomas et al. 2007). In contrast, several recent studies have suggested that the different effects of metabolic syndrome in older population. Higher blood pressure levels have been associated with better cognitive functioning and faster walking speed in elderly

adults (Odden et al. 2012; Zuccala et al. 2005). Faster walking speed, also often termed gait speed, has been shown to reflect muscle mass (Auyeung et al. 2014; Patil et al. 2013). Consistent with those studies, our finding suggest that higher  $ASM/Ht^2$  was associated with an increased risk of high blood pressure. In a more general sense, older age might represent a condition of frailty, which is associated with the epidemiological phenomenon of “reverse epidemiology” (Chien et al. 2012; Guder et al. 2015). In this perspective, our study supported that muscle mass is an independent risk factor for metabolic abnormalities. However, the aforementioned studies included hospitalized patients or, very old subjects or Western population, thus these findings are limited to apply to healthy older people.

Additionally, our findings for muscle mass may be explained by assuming that the higher muscle mass group includes subjects with both obesity and high fat mass. A study by Kimyagarov et al (2010), when body composition was analyzed according to the three BMI groups, subjects with normal BMI show a significantly increased absolute body fat and body fat/ $Ht^2$  , but not muscle mass from those in the low and high BMI groups (Kimyagarov et al. 2010). However, our study shows that increases in muscle mass have been shown to be related to increased body fat and grip strength. On the other hand, increases in body fat have been shown to be related to increased muscle mass but not grip strength. These finding are suggested that fat mass and muscle mass are not biologically independent. In our study, among the highest tertile of muscle mass they simultaneously included high body fat and high muscle mass groups, and low body fat and high muscle mass group, thus adjustment for body fat as a covariate might be inadequate.

In addition, we used ROC analysis to address the issue of discriminative performance. Body fat seems to be a better predictor of metabolic abnormalities in men, while muscle and fat mass indices are similar prediction in women. These result demonstrated that the pattern

and magnitude of body composition changes varied for the different indices of muscle and fat mass, was not similar for men and women (Strugnell et al. 2014).

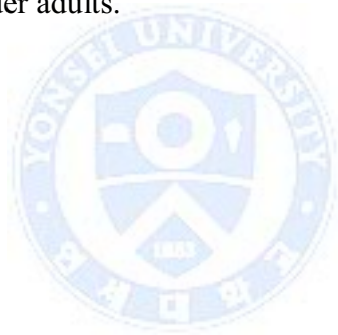
#### ***4. Limitations***

Our study has several limitations. First, muscle mass and fat mass does not directly assess the deposition of body composition such as DXA. Thus, we could not address the relationship between direct measures and metabolic abnormalities. Second, since the subjects were community-dwelling older adults, our findings may not be able to be generalized to older Korean adults from other racial/ethnic groups. Finally, our study was a cross-sectional analysis which did not establish a causative relationship between muscle mass and fat mass with metabolic abnormalities.



## V. CONCLUSIONS

The findings of the present study indicated that higher muscle mass and fat mass further increases the risks of metabolic abnormalities, such as high blood pressure, low HDL cholesterol, high glucose and metabolic syndrome even adjustment of age and body composition in older adult Korean women. This study adds to the growing knowledge on the better predictor of metabolic abnormalities is fat mass than muscle mass in men, and muscle mass is also predicted metabolic abnormalities in women. Further longitudinal studies are required to clarify the mechanism by which muscle mass is related to the development of metabolic abnormality among older adults.



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## ABSTRACT (KOREAN)

### ‘노년 인구의 근육량, 체지방량과 대사위험요인과의 관련성’

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#### 연구 배경 및 목적:

최근 노년 인구에서 sarcopenia는 대사이상과 관련성이 있다고 보고되고 있다. 그러나 sarcopenia의 기준은 통일되어 있지 않고 어떤 기준을 따라야 하는지에 대한 논의가 계속되고 있다. 이에 본 연구에서는 sarcopenia를 정의하기 이전에 근육의 절대량과 대사위험요인 간의 관련성을 분석하고자 하였다.

#### 연구 방법:

본 연구는 지역사회기반 전향적 코호트인 Korean Urban Rural Elderly (KURE) study의 일부로, 2014년에 연구 참여에 동의한 65세 이상의 성인을 대상으로 시행되었다. 대상자 중 917명은 체성분 검사와 혈액 검사 모두를 2014년에 시행하였으나 368명은 체성분 검사는 2014년에, 혈액 검사는 2012년에 시행하였다.

체성분은 인바디 720(바이오스페이스)를 통해 측정하였고, 대사위험요인 지표들은 공복 혈액에서 측정되었다. 근육량, 체지방량과 대사위험요인과의 관련성을 보기 위해 상관분석, 일반선형 및 다변량회귀 분석을 하였고 혼란변수로는 연령, 흡연 및 음주 습관, 신체활동 수준과 각각 근육량과 체지방량을 보정하였다.

## 연구 결과:

근육량을 3구간으로 나누어 보았을 때 남녀 모두에서 근육량이 증가할수록 체지방량도 통계적으로 유의하게 증가하였다. 체지방량과 모든 대사위험요인은 연령, 흡연 및 음주 습관, 신체활동 수준과 근육량을 보정하였을 때 남자와 여자 모두에서 통계적으로 유의한 관련성을 보였으나 근육량과 대사위험요인은 연령, 흡연 및 음주 습관, 신체활동 수준과 체지방량을 보정하였을 때 여자에서 근육량이 많을 수록 혈압이 높을 오즈비가 2.46 (95% CI 1.61-3.75), HDL이 낮을 오즈비가 2.25 (95% CI 1.49-3.38), 혈당이 높을 오즈비가 1.61 (95% CI 1.05-2.48)로 독립적인 관련성을 보였고, 남자에서는 HDL이 낮을 오즈비가 1.88 (95% CI 1.01-3.49)으로 나머지 위험요인과는 통계적으로 유의한 관련성을 보이지 않았다.

## 고찰:

본 연구에서는 노년 인구에서 체지방의 증가뿐 아니라 근육량의 증가도 대사위험요인과 높은 상관성을 보였고 성에 따라 결과에 차이가 있었다. 특히 남자에서는 근육량과 HDL이, 여자에서는 혈압, HDL, 혈당, 대사증후군과 독립적인 관련성이 있었다. 근육량, 체지방량과 대사위험요인과의 인과적인 관계에 대한 평가를 위해서는 추후 전향적인 연구가 뒷받침 되어야 할 것이다.