



Risk for metabolic syndrome in the population with visceral fat area measured by bioelectrical impedance analysis

Han Ho Jeon^{1,*}, Yong Kang Lee^{1,*}, Dong Hyun Kim¹, Haeyong Pak², Sang Yun Shin¹, and Jeong Hun Seo¹

¹Department of Internal Medicine,
²Institute of Health Insurance and
Clinical Research, National Health
Insurance Service Ilsan Hospital,
Goyang, Korea

Received: November 28, 2018
Revised : May 24, 2019
Accepted: June 2, 2019

Correspondence to
Jeong Hun Seo, M.D.

Department of Internal
Medicine, National Health
Insurance Service Ilsan Hospital,
100 Ilsan-ro, Ilsandong-gu,
Goyang 10444, Korea
Tel: +82-031-900-0974
Fax: +82-031-900-6967
E-mail: jhsuh@nhimc.or.kr

*These authors contributed
equally to this work.

Background/Aims: To investigate whether visceral fat area (VFA) measured by bioelectric impedance analysis (BIA) was associated with metabolic syndrome in subjects with and without obesity.

Methods: A total 23,202 participants who underwent medical check-ups were assessed. Participants were stratified by body mass index (BMI) and VFA. We evaluated six different groups for metabolic syndrome: Group 1 (normal weight and low VFA), Group 2 (normal weight and high VFA), Group 3 (overweight and low VFA), Group 4 (overweight and high VFA), Group 5 (obesity and low VFA), and Group 6 (obesity and high VFA).

Results: Metabolic syndrome traits and metabolic syndrome were significantly more prevalent in the high-VFA (≥ 100 cm²) subgroup in each BMI group. Adjusted logistic regression analyses revealed that the odds ratio for metabolic syndrome compared with Group 1 was the highest in Group 6 (24.53; 95% confidence interval [CI], 21.77 to 27.64). Notably, the odds ratio of Group 2 was higher than that of Group 3 (2.92; 95% CI, 2.30 to 3.69 vs. 2.57; 95% CI, 2.23 to 2.97).

Conclusions: Our study demonstrates that the combination of BMI assessment and VFA determination by BIA may be a useful method for predicting the risk of metabolic syndrome. The VFA by BIA may be a useful target for interventions to improve metabolic syndrome.

Keywords: Intra-abdominal fat; Metabolic syndrome; Electric impedance

INTRODUCTION

Due to recent changes in eating habits and lifestyle, the prevalence of obesity is increasing worldwide [1,2], regardless of race, age, and gender, and the incidence rates and complications of obesity-related diseases are increasing. People with obesity are more likely to develop metabolic disorders such as metabolic syndrome, type 2 diabetes, hypertension, dyslipidemia, ischemic heart disease, stroke, fatty liver, gallbladder disease, and thyroid disease [2-4].

Visceral fat plays an important role in the development of metabolic and cardiovascular diseases [5-7]. However, body mass index (BMI), the main measure used to determine obesity, does not account for the amount of muscle and fat, but simply accounts for body weight [8]. Therefore, there is a limit to predicting obesity-related disease. In addition, it is possible to evaluate abdominal obesity through the measurement of waist circumference [9], but there are limitations in that body fat percentage cannot be measured directly, and the reproducibility is poor due to large errors in the measurements

[10]. Methods for measuring visceral fat area (VFA) include dual energy X-ray absorptiometry, computed tomography (CT), and magnetic resonance imaging; however, these methods require expensive equipment and exposure to radiation, which limits their use. Current body composition analysis through bioelectrical impedance analysis (BIA) is the most widely used method because it has the advantage of measuring body fat and muscle mass easily and inexpensively [11-13].

In this study, we analyzed the relationship between metabolic syndrome in the obesity group defined by BMI and the obesity group defined by the VFA determined by bioelectrical impedance and assessed the risk of metabolic syndrome by subdividing the groups according to the combination of BMI and VFA.

METHODS

Study population

We identified 43,837 adults (≥ 18 years old) who underwent voluntary routine check-ups at the National Health Insurance Service Ilsan Hospital between January 2011 and December 2015. When participants underwent multiple examinations, we analyzed the data from their first visit. We excluded participants with BMI less than 18.5 kg/m². In total, 23,202 participants were included in this study. This study was approved by the Institutional Review Board of the National Health Insurance Service Ilsan Hospital (IRB No.: 2017-04-043-002). Written informed consent by the patients was waived due to a retrospective nature of our study.

Data collection

The participants arrived at the hospital after an overnight fast. Clinical and laboratory data were collected during the health examination. The height and weight of each participant were measured while he/she maintained a straight standing posture while wearing a light examination suit and no shoes, and the BMI was calculated. The waist circumference was measured mid-way between the lowest rib and the iliac crest. Visceral fat area was measured by a trained nurse using an InBody 720 (Biospace Co., Seoul, Korea) according to the manufacturer's protocol. The following clinical and laboratory data were collected: age, body weight (kg), body mass

index, smoking, drinking habits, waist circumference (cm), systolic blood pressure (mmHg), diastolic blood pressure (mmHg), uric acid (mg/dL), total cholesterol (mg/dL), high-density lipoprotein cholesterol (HDL-C; mg/dL), low-density lipoprotein cholesterol (LDL-C; mg/dL), triglycerides (mg/dL), fasting glucose (mg/dL), hemoglobin A_{1c} (HbA_{1c}, %), gamma-glutamyl transferase (GGT, U/L), aspartate aminotransferase (AST, U/L), and alanine aminotransferase (ALT, U/L).

Definitions

The cut-off point for obesity for Asians is defined by the World Health Organization as a BMI of 23.0 kg/m² [14]. The participants were divided into three groups by BMI: normal weight (18.5 to 22.9 kg/m²), overweight (23 to 24.9 kg/m²), and obesity (≥ 25 kg/m²). The definition of obesity by VFA was set as VFA ≥ 100 cm² [15].

Participants were categorized into six groups according to the combination of BMI and VFA. The six groups were (Fig. 1): (1) normal weight and VFA < 100 cm² (Group 1); (2) normal weight and VFA ≥ 100 cm² (Group 2); (3) overweight and VFA < 100 cm² (Group 3); (4) overweight and VFA ≥ 100 cm² (Group 4); (5) obesity and VFA < 100 cm² (Group 5); and (6) obesity and VFA ≥ 100 cm² (Group 6).

Metabolic syndrome was defined in accordance with the National Cholesterol Education Program Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults, Adult Treatment Panel III. Participants were defined as having metabolic syndrome if they had three or more of the following five factors: (1) waist circumference: male ≥ 90 cm, female ≥ 80 cm; (2) triglycerides: ≥ 150 mg/dL or use of medication; (3) HDL-C: male < 40 mg/dL, female < 50 mg/dL or use of medication; (4) blood pressure $\geq 130/85$ mmHg (systolic blood pressure ≥ 130 mmHg or diastolic blood pressure ≥ 85 mmHg) or use of medication; and (5) fasting glucose ≥ 100 mg/dL or use of medication.

Statistical analysis

Continuous variables are presented as mean \pm standard deviation. Categorical variables are presented as count (percentage). Differences between groups were tested by Student's *t* test for continuous variables and the chi-square test for categorical variables. Multiple logistic regression analysis was used to determine the risk of metabolic syndrome in each of the six groups. First, we used

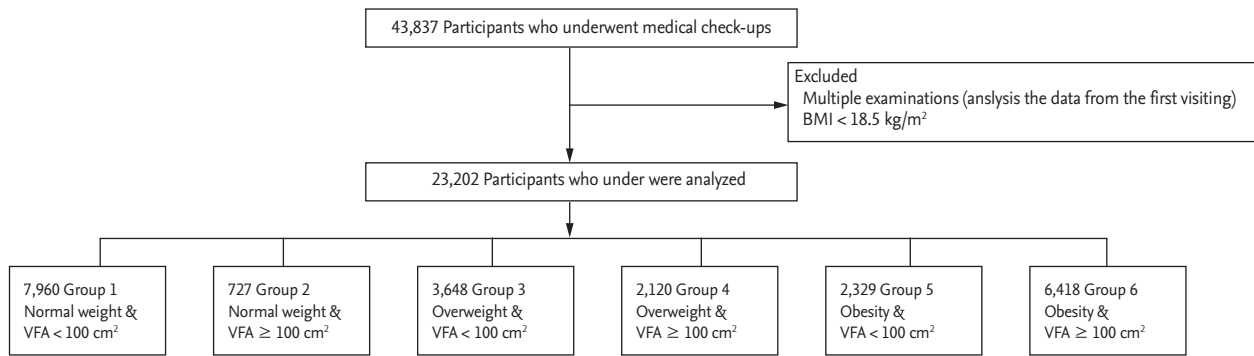


Figure 1. Study participant and categorization. Normal weight, body mass index (BMI) 18.5 to 22.9 kg/m²; overweight, BMI 23 to 24.9 kg/m²; obesity, BMI ≥ 25 kg/m². VFA, visceral fat area.

Table 1. Clinical characteristics of the study population stratified by gender

Characteristic	Male (n = 12,772)	Female (n = 10,430)	p value
Age, yr	47.9 ± 11.9	47.6 ± 12.5	0.028
Body weight, kg	73.5 ± 10.7	58.4 ± 8.3	< 0.001
BMI, kg/m ²	25.0 ± 3.0	23.3 ± 3.2	< 0.001
Waist circumference, cm	86.6 ± 7.8	80.6 ± 8.2	< 0.001
VFA, cm ²	106.5 ± 35.6	82.2 ± 31.3	< 0.001
Hypertension	14,02 (11.0)	674 (6.5)	< 0.001
Diabetes mellitus	2,229 (17.5)	1,111 (10.7)	< 0.001
Smoking ^a	9,666 (75.7)	745 (7.1)	< 0.001
Alcohol drinking ^b	4,842 (37.9)	3,298 (31.6)	< 0.001
Systolic BP, mmHg	123.7 ± 13.9	117.39 ± 15.1	< 0.001
Diastolic BP, mmHg	75.9 ± 10.6	71.49 ± 10.5	< 0.001
Total cholesterol, mg/dL	198.7 ± 37.5	195.8 ± 36.3	< 0.001
HDL-C, mg/dL	47.9 ± 11.4	56.67 ± 13.4	< 0.001
LDL-C, mg/dL	121.6 ± 33.2	116.85 ± 32.3	< 0.001
Triglyceride, mg/dL	153.2 ± 112.7	98.23 ± 65.5	< 0.001
HbA _{1c} , %	5.8 ± 0.8	5.7 ± 0.7	< 0.001
Metabolic syndrome	3,799 (29.7)	2,046 (19.6)	< 0.001
Fasting blood glucose, mg/dL	99.1 ± 23.5	94.1 ± 17.5	< 0.001
Uric acid, mg/dL	6.0 ± 1.2	4.4 ± 0.9	< 0.001
GGT, IU/L	39.8 ± 64.2	17.5 ± 15.7	< 0.001
AST, IU/L	27.5 ± 14.5	22.7 ± 9.9	< 0.001
ALT, IU/L	30.8 ± 23.5	20.0 ± 13.4	< 0.001

Values are presented as mean ± SD or number (%).

BMI, body mass index; VFA, visceral fat area; BP, blood pressure; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; HbA_{1c}, hemoglobin A_{1c}; GGT, gamma-glutamyl transferase; AST, aspartate transaminase; ALT, alanine transaminase.

^aSmoking: past smoker and current smoker.

^bAlcohol drinking: drink more than once a week.

Table 2. Correlation between VFA and other variables

Variable	Correlation coefficient
Age	0.209
BMI	0.701
Waist circumference	0.793
Systolic BP	0.349
Diastolic BP	0.307
Total cholesterol	0.148
HDL-C	-0.303
LDL-C	0.174
Triglyceride	0.322
Fasting glucose	0.229
HbA1c	0.228
Uric acid	0.340
GGT	0.166
AST	0.223
ALT	0.303

Values are the correlation coefficients and $p < 0.001$ for all analyses.

VFA, visceral fat area; BMI, body mass index; BP, blood pressure; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; HbA1c, hemoglobin A1c; GGT, gamma-glutamyl transferase; AST, aspartate transaminase; ALT, alanine transaminase.

the base model for the BMI and VFA at baseline. Model 1 was adjusted for age and sex, while Model 2 was adjusted for age, sex, diabetes mellitus, hypertension, alcohol intake, and smoking. The associations are presented as odds ratios (ORs) with 95% confidence intervals (CIs). All statistical analyses were performed with SAS 9.4 (SAS Institute, Cary, NC, USA). A p value below 0.05 was considered statistically significant.

RESULTS

Baseline characteristics of study population

The demographic and clinical characteristics of the participants are shown in Table 1. The mean age was 47.6 years for women and 47.9 years for men. The mean VFA was 82.2 cm² in women and 106.5 cm² in men. The prevalence of metabolic syndrome in women and men was 19.6% and 29.7% respectively. The mean BMI, blood pressure, uric acid, total cholesterol, LDL-C, triglycerides,

fasting blood glucose, GGT, AST, and ALT levels were significantly higher in men than in women. The mean HDL-C level was significantly higher in women than in men. Table 2 displays the correlations between VFA and various other parameters. The VFA correlated positively with BMI, blood pressure, uric acid, total cholesterol, LDL-C, triglycerides, fasting blood glucose, GGT, AST, and ALT levels, but negatively with HDL-C level.

Main characteristics of the groups defined by BMI and VFA

We divided participants into six groups according to the combination of BMI and VFA. The clinical characteristics of participants by gender according to BMI and VFA are shown in Tables 3 and 4. In men, the prevalence of metabolic syndrome was 6.1% in Group 1, 13.3% in Group 2, 13.1% in Group 3, 21.5% in Group 4, 25.4% in Group 5, and 56.2% in Group 6. In women, the prevalence of metabolic syndrome was 4.3% in Group 1, 21.6% in Group 2, 12.1% in Group 3, 30.9% in Group 4, 33.9% in Group 5 and 62.2% in Group 6. The subgroups of participants with VFA ≥ 100 cm² were generally older and had greater BMI and waist circumference than those with VFA < 100 cm². In addition, VFA ≥ 100 cm² was associated with the presence of hypertension, diabetes mellitus, metabolic syndrome, and metabolic parameters.

Presence of metabolic syndrome in groups defined by BMI and VFA

In variable-adjusted models, we estimated the risk of metabolic syndrome in each group defined by BMI and VFA (Table 5). In Model 1, the odds ratios for metabolic syndrome compared with Group 1 were 2.91 (95% CI, 2.31 to 3.67), 2.64 (95% CI, 2.29 to 3.04), 5.27 (95% CI, 4.55 to 6.11), 7.62 (95% CI, 6.63 to 8.75), and 25.44 (95% CI, 22.60 to 28.63) in Groups 2, 3, 4, 5, and 6, respectively. In Model 2, the respective odds ratios for metabolic syndrome compared with Group 1 were 2.92 (95% CI, 2.30 to 3.69), 2.57 (95% CI, 2.23 to 2.97), 5.16 (95% CI, 4.44 to 6.00), 7.45 (95% CI, 6.47 to 8.56), and 24.53 (95% CI, 21.77 to 27.64) in Groups 2, 3, 4, 5, and 6.

DISCUSSION

The following results were obtained through this study.

Table 3. Clinical characteristics of male according to BMI and VFA

Characteristic	Normal weight (n = 3,178)			Overweight (n = 3,548)			Obesity (n = 6,046)		
	VFA < 100 (n = 2,599)	VFA ≥ 100 (n = 579)	p value	VFA < 100 (n = 1,959)	VFA ≥ 100 (n = 1,589)	p value	VFA < 100 (n = 1,396)	VFA ≥ 100 (n = 4,650)	p value
Age, yr	47.3 ± 13.1	50.6 ± 12.2	< 0.001	47.3 ± 11.6	50.3 ± 11.6	< 0.001	47.1 ± 11.2	48.0 ± 11.2	0.009
Body weight, kg	62.5 ± 5.6	65.9 ± 5.5	< 0.001	69.5 ± 5.0	71.0 ± 5.4	< 0.001	76.0 ± 6.5	82.5 ± 9.7	< 0.001
BMI, kg/m ²	21.4 ± 1.1	22.1 ± 0.8	< 0.001	23.9 ± 0.6	24.1 ± 0.6	< 0.001	26.4 ± 1.3	27.8 ± 2.4	< 0.001
Hypertension	158 (6.1)	64 (11.1)	< 0.001	156 (8.0)	202 (12.7)	< 0.001	132 (9.5)	690 (14.8)	< 0.001
Diabetes mellitus	228 (8.8)	73 (12.6)	0.004	249 (12.7)	269 (16.9)	< 0.001	230 (16.5)	1180 (25.4)	< 0.001
Smoking ^a	1871 (72.0)	469 (81.0)	< 0.001	1380 (70.4)	1284 (80.8)	< 0.001	993 (71.1)	3669 (78.9)	< 0.001
Alcohol drinking ^b	836 (32.2)	209 (36.1)	< 0.001	706 (36.0)	608 (38.3)	0.173	531 (38.0)	1952 (42.0)	0.009
Waist circumference, cm	77.1 ± 3.4	81.6 ± 2.6	< 0.001	82.9 ± 2.3	85.7 ± 2.6	< 0.001	88.1 ± 3.4	94.0 ± 6.1	< 0.001
Metabolic syndrome	159 (6.1)	77 (13.3)	< 0.001	256 (13.1)	341 (21.5)	< 0.001	355 (25.4)	2611 (56.2)	< 0.001
Systolic BP, mmHg	118.1 ± 3.5	122.0 ± 13.3	< 0.001	121.3 ± 13.5	123.5 ± 13.2	< 0.001	124.4 ± 13.3	128.0 ± 13.3	< 0.001
Diastolic BP, mmHg	72.1 ± 10.1	74.9 ± 9.9	< 0.001	74.3 ± 10.4	75.8 ± 10.0	< 0.001	76.3 ± 10.4	78.8 ± 10.3	< 0.001
Total cholesterol, mg/dL	189.8 ± 34.7	199.7 ± 39.1	< 0.001	195.9 ± 36.2	204.2 ± 35.8	< 0.001	199.6 ± 35.9	203.2 ± 39.0	0.001
HDL-C, mg/dL	52.5 ± 13.0	49.6 ± 12.3	< 0.001	49.0 ± 11.2	48.3 ± 11.1	0.061	46.8 ± 10.8	45.0 ± 9.6	< 0.001
LDL-C, mg/dL	114.0 ± 30.9	122.0 ± 34.4	< 0.001	120.0 ± 31.2	126.4 ± 32.9	< 0.001	122.9 ± 32.9	124.8 ± 34.5	< 0.001
Triglyceride, mg/dL	108.9 ± 75.8	144.0 ± 105.0	< 0.001	131.9 ± 110.2	157.2 ± 104.0	< 0.001	151.2 ± 104.4	188.2 ± 125.7	< 0.001
HbA _{1c} , %	5.6 ± 0.8	5.8 ± 0.8	< 0.001	5.7 ± 0.8	5.8 ± 0.9	< 0.001	5.7 ± 0.8	5.9 ± 0.9	< 0.001
Fasting blood glucose, mg/dL	94.3 ± 22.8	98.1 ± 23.4	< 0.001	96.5 ± 23.8	100.5 ± 23.8	< 0.001	97.9 ± 20.0	103.0 ± 24.2	< 0.001
Uric acid, mg/dL	5.6 ± 1.1	5.8 ± 1.1	< 0.001	5.9 ± 1.1	6.0 ± 1.2	< 0.001	6.1 ± 1.2	6.3 ± 1.3	< 0.001
GGT, IU/L	34.1 ± 108.2	40.0 ± 77.0	0.157	32.0 ± 38.5	42.5 ± 47.6	< 0.001	35.7 ± 42.9	46.9 ± 45.6	< 0.001
AST, IU/L	25.0 ± 17.5	26.3 ± 14.1	0.049	25.3 ± 11.1	27.1 ± 11.6	< 0.001	27.0 ± 13.5	30.3 ± 14.8	< 0.001
ALT, IU/L	23.6 ± 28.5	27.4 ± 22.1	< 0.001	26.2 ± 15.2	30.1 ± 19.0	< 0.001	29.8 ± 17.9	37.8 ± 24.5	< 0.001

Values are presented as mean ± SD or number (%). Normal weight, BMI 18.5 kg/m²–22.9 kg/m²; Overweight, BMI 23 kg/m²–24.9 kg/m²; Obesity, BMI ≥ 25 kg/m². BMI, body mass index; VFA, visceral fat area; BP, blood pressure; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; HbA_{1c}, hemoglobin A_{1c}; GGT, gamma-glutamyl transferase; AST, aspartate transaminase; ALT, alanine transaminase.

^aSmoking: past smoker and current smoker.

^bAlcohol drinking: drink more than once a week.

Table 4. Clinical characteristics of female according to BMI and VFA

Characteristic	Normal weight (n = 5,509)		Overweight (n = 2,220)		Obesity (n = 2,701)	
	VFA < 100 (n = 5,361)	VFA ≥ 100 (n = 148)	VFA < 100 (n = 1,689)	VFA ≥ 100 (n = 531)	VFA < 100 (n = 933)	VFA ≥ 100 (n = 1,768)
Age, yr	43.8 ± 11.1	55.1 ± 11.5	48.6 ± 11.4	55.4 ± 12.2	49.1 ± 11.7	55.4 ± 12.4
Body weight, kg	53.3 ± 4.5	55.9 ± 4.5	59.3 ± 4.3	60.0 ± 4.8	64.8 ± 5.9	69.5 ± 8.8
BMI, kg/m ²	20.9 ± 1.2	22.0 ± 0.7	23.8 ± 0.6	24.1 ± 0.5	26.4 ± 1.4	28.3 ± 2.7
Hypertension	117 (2.2)	14 (9.5)	90 (5.3)	66 (12.4)	74 (7.9)	313 (17.7)
Diabetes mellitus	263 (4.9)	30 (20.3)	162 (9.6)	85 (16.0)	133 (14.3)	438 (24.8)
Smoking ^a	433 (8.1)	17 (11.5)	95 (5.6)	34 (6.4)	48 (5.1)	118 (6.7)
Alcohol drinking ^b	1,941 (36.2)	37 (25.0)	511 (30.3)	109 (20.5)	289 (31.0)	411 (23.2)
Waist circumference, cm	74.6 ± 3.5	80.0 ± 2.2	81.4 ± 2.2	84.2 ± 2.2	87.1 ± 3.7	93.3 ± 6.6
Metabolic syndrome	229 (4.3)	32 (21.6)	205 (12.1)	164 (30.9)	316 (33.9)	1,100 (62.2)
Systolic BP, mmHg	112.4 ± 13.5	122.0 ± 16.2	118.2 ± 14.2	122.9 ± 14.6	121.3 ± 14.7	127.9 ± 14.3
Diastolic BP, mmHg	68.8 ± 9.8	75.2 ± 11.2	71.8 ± 10.1	75.2 ± 10.0	73.2 ± 10.3	77.3 ± 9.9
Total cholesterol, mg/dL	190.7 ± 33.6	209.1 ± 38.5	197.3 ± 37.1	206.5 ± 38.2	199.2 ± 37.0	203.7 ± 38.9
HDL-C, mg/dL	59.9 ± 13.7	56.1 ± 13.1	55.5 ± 12.7	54.3 ± 11.8	53.3 ± 12.2	50.7 ± 11.7
LDL-C, mg/dL	110.6 ± 29.7	130.7 ± 34.0	119.5 ± 32.2	127.4 ± 34.6	121.7 ± 31.7	126.4 ± 35.2
Triglyceride, mg/dL	80.0 ± 47.3	108.6 ± 57.4	99.0 ± 62.8	119.5 ± 70.9	115.9 ± 79.2	135.7 ± 83.2
HbA _{1c} , %	5.5 ± 0.5	5.8 ± 1.0	5.6 ± 0.6	5.9 ± 0.9	5.8 ± 0.7	6.0 ± 0.9
Fasting blood glucose, mg/dL	90.0 ± 12.2	99.6 ± 32.3	93.5 ± 14.5	98.6 ± 18.9	96.8 ± 18.0	104.0 ± 25.2
Uric acid, mg/dL	4.2 ± 0.8	4.4 ± 1.0	4.3 ± 0.9	4.6 ± 1.0	4.6 ± 1.0	4.8 ± 1.0
GGT, IU/L	15.0 ± 13.1	20.6 ± 16.2	16.9 ± 12.4	19.3 ± 11.6	19.5 ± 16.8	23.7 ± 22.9
AST, IU/L	21.3 ± 8.0	23.0 ± 6.3	22.2 ± 8.1	23.9 ± 8.8	23.3 ± 8.9	26.9 ± 15.1
ALT, IU/L	17.2 ± 9.3	20.1 ± 7.7	19.7 ± 10.9	22.2 ± 11.6	21.9 ± 12.7	27.3 ± 21.8

Values are presented as mean ± SD or number (%). Normal weight, BMI 18.5–22.9 kg/m²; overweight, BMI 23–24.9 kg/m²; obesity, BMI ≥ 25 kg/m².

BMI, body mass index; VFA, visceral fat area; BP, blood pressure; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; HbA_{1c}, hemoglobin A_{1c}; GGT, gamma-glutamyl transferase; AST, aspartate transaminase; ALT, alanine transaminase.

^aSmoking: past smoker and current smoker.

^bAlcohol drinking: drink more than once a week.

Table 5. The odds ratio of metabolic syndrome in the group stratified by the combination of body mass index and visceral fat area

Variable	No.	Crude OR (95% CI)	p value	Model 1		Model 2	
				Adjusted OR (95% CI)	p value	Adjusted OR (95% CI)	p value
Group 1	7,960	1.00		1.00		1.00	
Group 2	727	3.44 (2.74–4.32)	< 0.001	2.91 (2.31–3.67)	< 0.001	2.92 (2.30–3.69)	< 0.001
Group 3	3,648	2.82 (2.45–3.25)	< 0.001	2.64 (2.29–3.04)	< 0.001	2.57 (2.23–2.97)	< 0.001
Group 4	2,120	6.10 (5.29–7.04)	< 0.001	5.27 (4.55–6.11)	< 0.001	5.16 (4.44–6.00)	< 0.001
Group 5	2,329	7.90 (6.89–9.05)	< 0.001	7.62 (6.63–8.75)	< 0.001	7.45 (6.47–8.56)	< 0.001
Group 6	6,418	26.75 (23.88–29.96)	< 0.001	25.44 (22.60–28.63)	< 0.001	24.53 (21.77–27.64)	< 0.001

Model 1. Adjust variable: age, sex; Model 2. Adjust variable: Model 1 + diabetes mellitus, hypertension, alcohol, smoking. OR, odds ratio; CI, confidence interval.

First, VFA estimated by BIA correlated well with waist circumference and metabolic syndrome components. Although previous studies have shown similar results regarding VFA [11,12,16,17], our study demonstrated the usefulness of VFA estimation by BIA in a large number of participants. Visceral fat is known to be associated with insulin resistance and diabetes [18]. Our study revealed that VFA was also associated with HbA_{1c} level. Additionally, we found that VFA was associated with the level of uric acid. Uric acid is not included in the diagnostic criteria for metabolic syndrome, but recent studies have revealed that increased uric acid level plays a causative role in the pathogenesis of metabolic, renal, and cardiovascular diseases and leads to increased mortality [19–21].

Second, when participants were divided into six groups, the odds ratio for metabolic syndrome tended to increase with group number, and the odds ratio for metabolic syndrome in Group 6 was very high at 24.53. Interestingly, when the normal-weight group was divided based on VFA, the odds ratio for metabolic syndrome in Group 2 (normal weight and VFA ≥ 100 cm²) was 2.92 compared with Group 1 (normal weight and VFA < 100 cm²). Additionally, the odds ratio for metabolic syndrome in Group 2 was higher than that in Group 3 (overweight and VFA < 100 cm²). The odds ratio for metabolic syndrome increased dramatically from 7.45 to 24.53 when participants with obesity based on BMI were further divided based on VFA (Group 5 vs. Group 6). These results suggest that VFA is an important component and cause of metabolic syndrome. Previous studies have demonstrated that VFA is associated with metabolic syndrome components and have proposed 100 cm² as a reasonable

cut-off point for VFA, suggesting that VFA < 100 cm² may be an important factor for improvement [15,22]. Visceral fat is associated with a greater cardiometabolic risk than subcutaneous fat or high BMI [23,24]. The adipocytokines induced by visceral fat are closely related with metabolic disorders [25], which develop into various metabolic and cardiovascular diseases. Visceral fat reduces the production of defensive adipocytokines such as adiponectin [25,26]. Changes in adiponectin in obesity play an important role in the development of metabolic and cardiovascular complications [25,27]. Circulating adiponectin level was found to correlate with VFA but not BMI in subjects with obesity [28].

The clinical significance of our study is as follows. The combination of BMI assessment and VFA determination by the BIA method allowed the diagnosis of individuals with an unrecognized but relatively high risk of metabolic syndrome (such as those with normal BMI and VFA greater than 100 cm²). In addition, our study confirmed that the risk of metabolic syndrome was significantly lower (OR, 7.45 vs. 24.53) when VFA was less than 100 cm² in participants with obesity based on BMI. In other words, VFA (cut-off point of 100 cm²) may be a useful target for interventions to improve metabolic syndrome.

The present study had several limitations. First, our study was limited by its cross-sectional and single-center retrospective design. Second, exercise as a habitual parameter was not evaluated in the present study because the data was not correctly acquired. However, the influence of this limitation was reduced by the large sample size in our study. Third, VFA was not measured by a standard method such as CT, which is the most ac-

curate and reliable method. However, VFA determined by BIA correlates well with that determined by CT [17,29].

In conclusion, this study demonstrated that the combination of BMI assessment and VFA determination by BIA may be a simple and useful method for predicting the risk of metabolic syndrome. The VFA by BIA may be a useful target for interventions to improve metabolic syndrome. Future interventional trials are needed to confirm the usefulness of targeting the VFA to improve metabolic syndrome.

KEY MESSAGE

1. Visceral fat area estimated by bioelectrical impedance analysis correlated well with waist circumference and metabolic syndrome components.
2. The combination of body mass index assessment and visceral fat area determination by the bioelectrical impedance analysis method allowed the diagnosis of individuals with an unrecognized but relatively high risk of metabolic syndrome.
3. Visceral fat area (cut-off point of 100 cm²) by bioelectrical impedance analysis may be a useful target for interventions to improve metabolic syndrome.

Conflict of interest

No potential conflict of interest relevant to this article was reported.

Acknowledgments

This study was supported by a grant from the National Health Insurance Service Ilsan Hospital, Republic of Korea. The authors acknowledge the effort of the health screening group at the National Health Insurance Service Ilsan Hospital, Republic of Korea.

REFERENCES

1. Finucane MM, Stevens GA, Cowan MJ, et al. National, regional, and global trends in body-mass index since 1980: systematic analysis of health examination surveys and epidemiological studies with 960 country-years and 9.1 million participants. *Lancet* 2011;377:557-567.
2. Bauer UE, Briss PA, Goodman RA, Bowman BA. Prevention of chronic disease in the 21st century: elimination of the leading preventable causes of premature death and disability in the USA. *Lancet* 2014;384:45-52.
3. Schienkiewitz A, Mensink GB, Scheidt-Nave C. Comorbidity of overweight and obesity in a nationally representative sample of German adults aged 18-79 years. *BMC Public Health* 2012;12:658.
4. Calle EE, Rodriguez C, Walker-Thurmond K, Thun MJ. Overweight, obesity, and mortality from cancer in a prospectively studied cohort of U.S. adults. *N Engl J Med* 2003;348:1625-1638.
5. Matsuzawa Y, Funahashi T, Nakamura T. The concept of metabolic syndrome: contribution of visceral fat accumulation and its molecular mechanism. *J Atheroscler Thromb* 2011;18:629-639.
6. Fox CS, Massaro JM, Hoffmann U, et al. Abdominal visceral and subcutaneous adipose tissue compartments: association with metabolic risk factors in the Framingham Heart Study. *Circulation* 2007;116:39-48.
7. Albu JB, Kovera AJ, Johnson JA. Fat distribution and health in obesity. *Ann N Y Acad Sci* 2000;904:491-501.
8. Gallagher D, Visser M, Sepulveda D, Pierson RN, Harris T, Heymsfield SB. How useful is body mass index for comparison of body fatness across age, sex, and ethnic groups? *Am J Epidemiol* 1996;143:228-239.
9. Carr DB, Utzschneider KM, Hull RL, et al. Intra-abdominal fat is a major determinant of the National Cholesterol Education Program Adult Treatment Panel III criteria for the metabolic syndrome. *Diabetes* 2004;53:2087-2094.
10. Dhaliwal SS, Welborn TA. Measurement error and ethnic comparisons of measures of abdominal obesity. *Prev Med* 2009;49:148-152.
11. Ranasinghe C, Gamage P, Katulanda P, Andraweera N, Thilakarathne S, Tharanga P. Relationship between body mass index (BMI) and body fat percentage, estimated by bioelectrical impedance, in a group of Sri Lankan adults: a cross sectional study. *BMC Public Health* 2013;13:797.
12. Unno M, Furusyo N, Mukae H, Koga T, Eiraku K, Hayashi J. The utility of visceral fat level by bioelectrical impedance analysis in the screening of metabolic syndrome: the results of the Kyushu and Okinawa Population Study (KOPS). *J Atheroscler Thromb* 2012;19:462-470.
13. Kushner RF, Gudivaka R, Schoeller DA. Clinical char-

- acteristics influencing bioelectrical impedance analysis measurements. *Am J Clin Nutr* 1996;64:423S-427S.
14. WHO Expert Consultation. Appropriate body-mass index for Asian populations and its implications for policy and intervention strategies. *Lancet* 2004;363:157-163.
 15. Examination Committee of Criteria for 'Obesity Disease' in Japan; Japan Society for the Study of Obesity. New criteria for 'obesity disease' in Japan. *Circ J* 2002;66:987-992.
 16. Lewin A, Pannier B, Meline J, Karusisi N, Thomas F, Chaix B. Residential neighborhood, geographic work environment, and work economic sector: associations with body fat measured by bioelectrical impedance in the RECORD Study. *Ann Epidemiol* 2014;24:180-186.
 17. Sakamaki K, Maejima Y, Tokita Y, et al. Impact of the visceral fat area measured by dual impedance method on the diagnostic components of metabolic diseases in a middle-aged Japanese population. *Intern Med* 2016;55:1691-1696.
 18. Koh H, Hayashi T, Sato KK, et al. Visceral adiposity, not abdominal subcutaneous fat area, is associated with high blood pressure in Japanese men: the Ohtori study. *Hypertens Res* 2011;34:565-572.
 19. Kang DH, Park SK, Lee IK, Johnson RJ. Uric acid-induced C-reactive protein expression: implication on cell proliferation and nitric oxide production of human vascular cells. *J Am Soc Nephrol* 2005;16:3553-3562.
 20. Nakagawa T, Mazzali M, Kang DH, et al. Hyperuricemia causes glomerular hypertrophy in the rat. *Am J Nephrol* 2003;23:2-7.
 21. Hakoda M, Masunari N, Yamada M, et al. Serum uric acid concentration as a risk factor for cardiovascular mortality: a longterm cohort study of atomic bomb survivors. *J Rheumatol* 2005;32:906-912.
 22. Tatsumi Y, Nakao YM, Masuda I, et al. Risk for metabolic diseases in normal weight individuals with visceral fat accumulation: a cross-sectional study in Japan. *BMJ Open* 2017;7:e013831.
 23. Despres JP, Moorjani S, Lupien PJ, Tremblay A, Nadeau A, Bouchard C. Regional distribution of body fat, plasma lipoproteins, and cardiovascular disease. *Arteriosclerosis* 1990;10:497-511.
 24. Shah RV, Murthy VL, Abbasi SA, et al. Visceral adiposity and the risk of metabolic syndrome across body mass index: the MESA Study. *JACC Cardiovasc Imaging* 2014;7:1221-1235.
 25. Kishida K, Funahashi T, Shimomura I. Adiponectin as a routine clinical biomarker. *Best Pract Res Clin Endocrinol Metab* 2014;28:119-130.
 26. Weyer C, Funahashi T, Tanaka S, et al. Hypoadiponectinemia in obesity and type 2 diabetes: close association with insulin resistance and hyperinsulinemia. *J Clin Endocrinol Metab* 2001;86:1930-1935.
 27. Santaniemi M, Kesaniemi YA, Ukkola O. Low plasma adiponectin concentration is an indicator of the metabolic syndrome. *Eur J Endocrinol* 2006;155:745-750.
 28. Kishida K, Kim KK, Funahashi T, Matsuzawa Y, Kang HC, Shimomura I. Relationships between circulating adiponectin levels and fat distribution in obese subjects. *J Atheroscler Thromb* 2011;18:592-595.
 29. Ogawa H, Fujitani K, Tsujinaka T, et al. InBody 720 as a new method of evaluating visceral obesity. *Hepatogastroenterology* 2011;58:42-44.