Weight Gain Predicts Metabolic Syndrome among North Korean Refugees in South Korea

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Abstract: Previous cross-sectional studies showed that immigrants from low-income to high-income countries have higher risk of cardiovascular disease and type 2 diabetes mellitus. We investigated the association between weight gain during the resettlement in South Korea and risk of metabolic syndrome (MetS) among North Korean refugees (NKRs) in this cross-sectional study. In total, 932 NKRs aged 20–80 years in South Korea voluntarily underwent health examination from 2008 to 2017. We compared the risk of MetS and its components between the weight gain group (gained ≥5 kg) and the non-weight gain group (gained <5 kg, maintained or lost body weight) during resettlement in South Korea after defection from North Korea. Multiple logistic regression analysis predicted odds ratio of MetS on the basis of weight change, adjusting for covariates and current body mass index (BMI). We also evaluated the difference in body composition of NKRs between two groups. The prevalence of MetS in the weight gain group was 26%, compared to 10% in the non-weight gain group (p-value < 0.001). The weight gain group had a two-fold higher risk of MetS than the non-weight gain group after adjusting for current BMI (odds ratio 1.875, p-value = 0.045). The prevalence of central obesity, impaired fasting glucose, elevated blood pressure, and hypertriglyceridemia were higher in the weight gain group than the non-weight gain group (36% vs. 12%, p-value < 0.001; 32% vs. 19%, p-value < 0.001; 34 vs. 25%, p-value = 0.008; 19% vs. 13%, p-value = 0.025, respectively). The analysis of body composition showed that the percentage of body fat in the weight gain group was higher than in the non-weight gain group, indicating increased fat mass rather than muscle mass in the weight gain group as their body weight increased during resettlement (33.4 ± 6.5% vs. 28.88 ± 7.40%, p < 0.005). Excess weight gain after defection from North Korea increased the risk of MetS among NKRs in South Korea. It is necessary to monitor weight change among NKRs and their effect on their metabolic health in the long term.

Keywords: North Korean refugee; weight gain; metabolic syndrome; percentage body fat

1. Introduction

As the prevalence of obesity has increased worldwide due to dietary change and sedentary lifestyles, the prevalence of metabolic syndrome (MetS) has also rapidly increased [1]. MetS could be a public health problem in that it could be a risk factor for diabetes mellitus and cardiovascular disease [2,3]. Obesity is a well-known risk factor for MetS in the general adult population [4–6]. Obesity is etiologically linked with chronic low-grade inflammatory conditions that contribute to the pathogenesis of metabolic dysfunction and MetS [7]. Previous studies have shown that weight gain had a significant
relationship with MetS [8,9]. According to a previous prospective study, 4.5 kg of weight gain during a 13-year follow-up increased the risk of MetS by 23% [8]. Another cross-sectional study revealed that the changes of body mass index (BMI) in early adulthood (25–40 years) were strongly associated with increased risk of metabolic components, such as increased Hemoglobin A1c (Hb A1c) or low high-density lipoprotein (HDL) cholesterol level [9]. Cross-sectional studies among Japanese populations revealed that weight gain increased the risk of MetS, even in a non-obese population [10,11]. Hashimoto et al. showed that weight gain after age of 20 years increased the risk of MetS by twofold in non-obese individuals [10].

Immigrants from low-income countries experienced weight gain when they migrated to high-income countries due to westernized lifestyle and dietary changes [12,13]. Immigrant women from Turkey and Chile to Sweden had a higher risk of obesity due to change of dietary habits and low physical activity in a cross-sectional study [12]. Another longitudinal study among immigrant Chinese women in the United States (US) showed that increasing length of US residence was associated with increased energy density in diet and sugar intake [13]. Furthermore, previous studies showed that immigrants from low- to high-income countries had a higher risk of cardiovascular disease and type 2 diabetes mellitus than the host population [14,15]. A cross-sectional study of South Asian immigrants in the US showed that South Asians had coronary artery disease more commonly than Caucasians [15]. However, few studies have assessed how this weight change among immigrants affects their metabolic health during resettlement in host country [16].

The ethnic background of the population of the Korean Peninsula remained unchanged for 5000 years, but Korea was divided into North and South Korea after the Korean War. While South Korea has achieved rapid economic growth by introducing a free-market economy system, North Korea has experienced severe poverty due to collapse of communism and continuous natural disasters. According to World Health Organization (WHO, Geneva, Switzerland) survey data in 2008, the North Korean population aged 25–64 years had very low obesity prevalence, 4.2% in men and 4.6% in women. On the contrary, obesity prevalence in the same age group is 38.9% in South Korean men and 25.3% in women according to a 2008 Korean National Health and Nutrition Examination Survey [17,18]. Exhausted from poverty and hunger, many North Koreans have left North Korea and resettled in South Korea through transit countries [19,20]. The number of North Korean refugees (NKRs) in South Korea reached 33,000 in December 2019 [21]. We previously showed that NKRs experienced weight gain and approached the average BMI of South Korean population 10 years after defection from North Korea [22]. In addition, while NKRs still had a lower BMI than the South Korean population, the prevalence of MetS among NKRs was similar with that of the South Korean population [23].

In the present study, we investigated the association between weight gain and the risk of MetS among NKRs in South Korea. NKRs with more than three of the five components were classified as having MetS (central obesity, impaired fasting glucose, elevated blood pressure, hypertriglyceridemia, and low HDL cholesterol). We hypothesized that NKRs who gained weight more than 5 kg would have a higher risk of MetS. We also examined the difference in body composition between the two groups of NKRs with and without 5 kg or more weight gain after defection from North Korea by bioelectrical impedance analysis (BIA).

2. Methods

2.1. Study Participants

This study was conducted in the phase 1 survey of the North Korean Refugee Health in South Korea (NORNS) study [24]. The NORNS study is composed of two phases. The phase 1 study is a cross-sectional survey, and the phase 2 study is a follow-up survey 3.5 years after the phase 1 study. The NORNS study has taken place at Korea University Anam Hospital since October 2008 and still ongoing. Hana center, a representative welfare center supported by the South Korean government for assisting NKRs’ resettlement in South Korea, posted a
notice about the study on the Internet and by telephone once a month, and NKRs who have expressed their intention to participate can take part in the study. From October 2008 to March 2017, a total of 932 NKRs (men = 192, women = 740) above 20 years living in Seoul, South Korea, were recruited. The participant’s survey consisted of health questionnaire and medical examination. Medical examination comprised anthropometric assessment, blood pressure, biochemical measurement, and analysis of body composition through BIA. We have described the protocols and methods of the study in detail previously [24]. Of the 932 participants, we excluded those whom data of body weight, waist circumference (WC), or history of body weight were missing or unreliable. Finally, 799 NKRs (men = 162, women = 637) aged 20–80 years were analyzed. All participants provided written informed consent and the study was approved by the Institutional Review Board of Korea University Anam Hospital (approval number: Ed08023).

2.2. Measurements

2.2.1. Sociodemographic Characteristics

We produced a 42-item health questionnaire based on the existing questionnaire of Korea National Health and Nutrition Examination Survey (KNHANES) that is composed of six domains (demographic and migration information, disease history, mental health status, health-related lifestyles, female reproductive health, and sociocultural adaptation) (Supplementary Material, Table S1). In this study, we used the sociodemographic information including age, sex, health-related lifestyle factors, education level, occupation, current income in South Korea, and migration information (time of defection from North Korea, time of arrival in transit country and South Korea). Health-related lifestyle factors included smoking, frequent alcohol drinking (more than one bottle of alcohol per week), and regular exercise (vigorous physical activity more than 1 h per week). Higher income was defined as more than 10^6 Korean won (KRW)/month, and higher education level was defined as above college graduate. Participants were asked to recall their body weights in North Korea, transit countries, and South Korea. For the completeness of the questionnaire, we had the NKR interview with a medical doctor who defected from North Korea.

2.2.2. Anthropometric Measurement

The health examinations were conducted by licensed nurses and trained volunteer. Body weight was measured in light clothing and height was measured without shoes by automatic system (GL-150; G-Tech International, Seoul, South Korea). BMI was calculated as weight (kg) divided by squared height (m^2). WC was measured at minimum abdominal girth, midpoint between the lowest rib and iliac crest by nurses [25]. Systolic and diastolic blood pressure were measured by an autonomic blood pressure monitor (TM-2655P; Biospace, Tokyo, Japan) on the arm of a seated participant who rested in a sitting position for 10 min before measurement and used the mean value in the analysis [26]. Participants underwent BIA (Inbody720, Biospace, Seoul, South Korea) while wearing light clothes to analyze body muscle mass, fat mass, and percentage of body fat.

2.2.3. Biochemical Measurement

Blood samples were drawn in the morning after an overnight fasting. Serum total cholesterol, triglyceride, HDL cholesterol, low density lipoprotein (LDL) cholesterol, and liver enzyme were measured using enzymatic method (TBA 200-FR; Toshiba, Tokyo, Japan). Plasma glucose concentrations were measured by the glucose oxidase method. Serum insulin levels were measured by radioimmunoassay (Diasource, Nivelles, Belgium). The homeostasis model assessment–insulin resistance (HOMA–IR) was computed using following formula: fasting plasma glucose (mg/dL) × fasting serum insulin (µU/mL)/405 [27]. Insulin secretary function (HOMA–ß) was calculated with following formula: 360 × fasting insulin (µU/mL)/fasting glucose (mg/dL)–63 [27].

Metabolic syndrome was defined when more than three of the following criteria were met according to modified NCEP–ATP III guideline [28]: (1) WC ≥ 90 cm (men), ≥85 cm
(women); (2) fasting glucose ≥ 100 mg/dL; (3) systolic blood pressure ≥ 130 mmHg or diastolic blood pressure ≥ 85 mmHg; (4) HDL cholesterol < 40 mg/dL (men), <50 mg/dL (women); (5) triglycerides ≥ 150 mg/dL. Participants who took medication for diabetes or hypertension were regarded as having met the criteria for elevated fasting glucose or elevated blood pressure.

2.2.4. Classification of NKRs by Change in Weight after Defection from North Korea

To investigate the effect of weight gain after defection from North Korea on the risk of MetS, we classified NKRs into weight gain group (gained body weight more than 5 kg after defection from North Korea) and non-weight gain group (gained less than 5 kg, maintained or lost their body weight during resettlement in South Korea), and compared metabolic health between two groups. The cut-off value of 5 kg was chosen on the basis of previous large prospective cohort studies that investigated the association between weight change and coronary heart disease [29,30].

2.3. Statistical Analysis

Comparison of sociodemographic and metabolic characteristics in NKRs were presented by weight gain and non-weight gain group. Comparison of categorical variables were described as frequency (percentage) with Pearson $\chi^2$ test. Continuous variables were described as mean ± standard deviation using analysis of variance (ANOVA). Relationship between weight gain and risk of MetS was investigated with multivariate logistic regression analyses using MetS risk as a dependent variable. Odds ratio (OR) and 95% confidence intervals (95% CI) were calculated with non-weight gain group as the reference group. The logistic regression was conducted in two steps. In Model 1, the ORs were adjusted for age, sex, smoking, alcohol drinking, regular exercise, education, income, stay duration of transit countries, and defection period. In the second step (Model 2), the ORs were adjusted by adding the current BMI and lipid medication to covariates in Model 1. p-value < 0.05 was considered as significant. All statistical analyses were performed using SPSS 26.0 software (SPSS Inc., Chicago, IL, USA).

3. Results

3.1. Characteristics of The Study Population

The demographic characteristics of the NKRs according to weight gain after defection from North Korea are shown in Table 1. Among 799 participants, 253 (32%) were assigned to the weight gain group and 546 (68%) of NKRs to the non-weight gain group. The mean age and sex distribution were similar between the two groups. The weight gain group had higher mean value of body weight, BMI, and WC than the non-weight gain group. Mean fasting glucose, triglyceride, ALT levels, and blood pressure were higher in the weight gain group than non-weight gain group. Mean value of HOMA–IR was higher in weight gain group than non-weight gain group (1.68 ± 1.11 vs. 1.38 ± 1.2, p-value = 0.002). There was no significant difference in alcohol drinking, smoking, and physical activity between the two groups. Defection period from North Korea was longer in the weight gain group than non-weight gain group (7.91 ± 4.97 vs. 6.67 ± 4.62 years, p-value = 0.001). The weight gain group had a lower mean body weight and BMI in North Korea than the non-weight gain group, although they had a higher BMI and body weight than non-weight gain group on examination in South Korea. The mean weight change was 10 ± 4.8 kg in weight gain group and −0.5 ± 3.96 kg in non-weight gain group (p-value < 0.001).

3.2. Prevalence of Individual Metabolic Components in The Weight Gain and Non-Weight Gain Groups

Metabolic syndrome was 2.6 times more prevalent in the weight gain group than the non-weight gain group (26% vs. 10%, p-value < 0.001) (Table 2). The prevalence of central obesity in the weight gain group was three times higher than in non-weight gain group (36% vs. 12%, p-value < 0.001). Prevalence of impaired fasting glucose and
Elevated blood pressure were higher in the weight gain group than in the non-weight gain group (32% vs. 19%, $p$-value < 0.001, and 34% vs. 25%, $p$-value = 0.008, respectively). Hypertriglyceridemia was also more prevalent in the weight gain group than in the non-weight gain group (19% vs. 13%, $p$-value = 0.025). The proportion of subjects with low HDL cholesterol showed no difference between the weight gain and non-weight gain groups (38% vs. 36%, $p$-value = 0.434).

| Table 1. Characteristics of North Korean refugees according to degree of weight gain. |
| --- |
| **Weigh Gain < 5 kg (n = 546)** | **Weight Gain ≥ 5 kg (n = 253)** | **p-Value** |
| Age | 43.25 ± 12.31 | 44.78 ± 12.97 | 0.108 |
| Sex (male) | 109/545 (20%) | 53/253 (21%) | 0.794 |
| Height (cm) | 156.26 ± 7.28 | 157.3 ± 7.41 | 0.061 |
| Body weight (kg) | 53.60 ± 7.37 | 60.7 ± 6.85 | <0.001 |
| BMI (kg/m$^2$) | 21.98 ± 2.47 | 24.48 ± 2.68 | <0.001 |
| Waist circumference (cm) | 76.50 ± 7.79 | 83.69 ± 8.57 | <0.001 |
| Fasting glucose (mg/dL) | 93.21 ± 10.76 | 95.93 ± 13.64 | 0.003 |
| Total cholesterol (mg/dL) | 174.93 ± 37.21 | 178.85 ± 38.60 | <0.001 |
| Triglyceride (mg/dL) | 97.61 ± 92.92 | 103.04 ± 78.62 | 0.023 |
| HDL cholesterol (mg/dL) | 54.86 ± 28.58 | 51.38 ± 12.16 | 0.065 |
| AST (mg/dL) | 21.42 ± 11.45 | 22.22 ± 14.73 | 0.407 |
| ALT (mg/dL) | 18.01 ± 17.65 | 21.75 ± 23.08 | 0.012 |
| HOMA-IR | 1.38 ± 1.29 | 1.68 ± 1.11 | 0.002 |
| HOMA-ß | 74.06 ± 45.76 | 79.67 ± 44.90 | 0.112 |
| BMI in North Korea (kg/m$^2$) | 22.14 ± 2.69 | 20.46 ± 2.71 | <0.001 |
| Weight in North Korea (kg) | 54.10 ± 7.78 | 50.74 ± 8.32 | <0.001 |
| Weight change (kg) | −0.5 ± 3.96 | 10.0 ± 4.8 | <0.001 |
| Defection period (yr) | 6.67 ± 4.62 | 7.91 ± 4.97 | 0.001 |
| Stay in transit country (yr) | 3.58 ± 3.76 | 3.86 ± 3.68 | 0.333 |
| Income (>10$^6$ KRW/month) | 95.49 ± 93.73 | 108.15 ± 132.28 | 0.196 |
| Higher education | 115/546 (21%) | 48/253 (19%) | 0.582 |

Categorical variables are shown as number (%); continuous variables are given as mean ± standard deviation. * Frequent alcohol drinking*: more than one bottle of alcohol per week. † Regular exercise: vigorous activity more than one hour per week. ‡ Higher education: more than college graduate. Abbreviations: BMI, body mass index; HDL cholesterol, high density lipoprotein cholesterol; HOMA-IR, homeostatic model assessment for insulin resistance; HOMA-ß, homeostatic model assessment for beta cell function; BP, blood pressure; KRW, Korean won.

| Table 2. Comparison of metabolic syndrome and its components according to degree of weight gain. |
| --- |
| **Weigh Gain < 5 kg (n = 543)** | **Weight Gain ≥ 5 kg (n = 250)** | **p-Value** |
| Metabolic syndrome | 53/527 (10%) | 63/243 (26%) | <0.001 |
| Central obesity * | 64/537 (12%) | 89/247 (36%) | <0.001 |
| Impaired fasting glucose † | 103/543 (19%) | 80/250 (32%) | <0.001 |
| Elevated blood pressure ‡ | 134/538 (25%) | 85/250 (34%) | 0.008 |
| Hypertriglyceridemia § | 72/543 (13%) | 47/250 (19%) | 0.025 |
| Low HDL cholesterol † | 195/543 (36%) | 95/250 (38%) | 0.434 |

* Central obesity, ≥90 cm (men), ≥85 cm (women); † impaired fasting glucose, fasting glucose ≥100mg/dL; ‡ elevated blood pressure, systolic BP ≥130mmHg or diastolic BP ≥ 85 mmHg; § hypertriglyceridemia, triglyceride ≥150 mg/dL; †† low HDL cholesterol <40mg/dL (men), <50 mg/dL (women).

3.3. Number of Metabolic Components in Weight Gain and Non-Weight Gain Groups

The weight gain group had a higher proportion of subjects with three or more metabolic components than the non-weight gain group (26% vs. 10%, $p < 0.001$) (Figure 1).
The non-weight gain group had a higher proportion of subjects with no or one metabolic component than the weight gain group (72% vs. 54%, \( p < 0.001 \)).

![Figure 1](image)

**Figure 1.** Number of metabolic syndrome components according to degree of weight gain.

### 3.4. Body Composition in Weight Gain and Non-Weight Gain Groups

In analyzing body composition between the weight gain group and non-weight gain group by BIA, we found that the difference of muscle mass was not large (21.88 ± 4.60 kg vs. 20.7 ± 4.11 kg, \( p \)-value < 0.001) (Table 3). The weight gain group had a higher mean body fat mass and percentage of body fat than the non-weight gain group (20.32 ± 5.01 kg vs. 15.73 ± 5.88 kg, \( p \)-value < 0.001, 33.4 ± 6.53% vs. 28.88 ± 7.40%, \( p \)-value < 0.001, respectively).

| Variables               | Weight Gain < 5 kg (n = 545) | Weight Gain ≥ 5 kg (n = 252) | \( p \)-Value |
|-------------------------|-------------------------------|-------------------------------|--------------|
| Body weight (kg)        | 53.60 ± 7.37                  | 60.7 ± 8.65                   | <0.001       |
| Muscle mass (kg)        | 20.67 ± 4.11                  | 21.88 ± 4.60                  | <0.001       |
| Fat mass (kg)           | 15.73 ± 5.88                  | 20.32 ± 5.01                  | <0.001       |
| Percentage of body fat (%) | 28.88 ± 7.40               | 33.4 ± 6.53                   | <0.001       |

Variables are given as mean ± standard deviation.

### 3.5. Risk Factors of Metabolic Syndrome among NKRs

Table 4 shows the result of multiple logistic regression analyses to evaluate the risk of MetS among the weight gain group when compared to non-weight gain group. The weight gain group had a twofold higher risk of MetS than the non-weight gain group after adjusting for multiple sociodemographic variables and current BMI (OR, 1.875; 95% CI, 1.013–3.468). Old age, female sex, and BMI were positively associated with MetS in multivariate logistic regression analysis. However, frequent alcohol drinking, current smoking, regular exercise, higher education, higher income, stay duration in transit countries, deflection period, and lipid medication had no significant association with risk of MetS.
Table 4. Odds ratio of metabolic syndrome risk among North Korean refugees.

|                              | Model 1 (n = 769) | p-Value | Model 2 (n = 509) | p-Value |
|------------------------------|-------------------|---------|-------------------|---------|
| OR (95% CI)                  | OR (95% CI)       |         |                   |         |
| Age *                        | 1.073             | < 0.001 | 1.056             | < 0.001 |
|                              | (1.049–1.097)     |         | (1.030–1.081)     |         |
| Sex (male as reference)      | 2.708             | 0.032   | 2.852             | 0.029   |
| Weight gain (≥5 kg)          | 3.392             | < 0.001 | 1.875             | 0.045   |
|                              | (1.973–5.832)     |         | (1.013–3.468)     |         |
| Current smoker               | 2.479             | 0.100   | 3.128             | 0.051   |
| (nonsmoker as reference)     | (0.841–7.307)     |         | (0.997–9.820)     |         |
| Frequent alcohol drinking †  | 0.962             | 0.887   | 0.937             | 0.827   |
| (nondrinker as reference)    | (0.560–1.62)      |         | (0.520–1.686)     |         |
| Regular exercise ‡           | 0.818             | 0.468   | 0.724             | 0.290   |
|                              | (0.476–1.407)     |         | (0.398–1.317)     |         |
| Stay in transit country (years) | 0.973         | 0.629   | 0.988             | 0.853   |
|                              | (0.870–1.088)     |         | (0.873–1.119)     |         |
| Defection period (years)     | 1.010             | 0.817   | 0.979             | 0.644   |
|                              | (0.930–1.096)     |         | (0.893–1.072)     |         |
| Higher education ‡           | 0.843             | 0.612   | 0.728             | 0.393   |
|                              | (0.435–1.633)     |         | (0.351–1.509)     |         |
| Higher income (>$10^6$ KRW/month) | 0.579      | 0.094   | 0.634             | 0.190   |
| BMI                          | (0.305–1.098)     |         | (0.321–1.254)     |         |
|                              |                   |         | 1.312             | 0.001   |
|                              |                   |         | (1.177–1.463)     |         |
| Lipid medication             | 0.866             | 0.480   |                   |         |
|                              | (0.651–0.125)     |         |                   |         |

* Age as continuous variable; † frequent alcohol drinking: more than one bottle of alcohol per week; ‡ regular exercise: vigorous activity more than one hour per week; § higher education: more than college graduate; abbreviations: OR, odds ratio; CI, confidence interval; BMI, body mass index; KRW, Korean won. Model 1, adjusted by age, sex, current smoking, alcohol drinking, regular exercise, stay period in transit countries, defection period, education, income. Model 2, adjusted by Model 1, lipid medication, and BMI.

4. Discussion

We investigated the association between weight gain of more than 5 kg and the risk of MetS among NKR in South Korea. More than 5 kg weight gain after defection from North Korea was associated with a twofold higher risk of MetS after adjusting for BMI. A previous prospective study has reported a strong relationship between weight gain and the risk of MetS in Caucasian and African American populations (MetS defined according to ATP III guideline) [8]. A cross-sectional study of the Japanese adult population revealed that not only obese but also non-obese individuals who gained weight more than 10 kg over 20 years had a fivefold higher risk of MetS (defined as abdominal obesity plus one of three risk factors, high blood pressure, hypertriglyceridemia or low HDL cholesterol, high fasting glucose) [11]. In addition to long-term weight gain, the prospective study has suggested that 1 year weight gain also increases the risk of metabolic components such as abdominal circumference, blood pressure, and serum triglyceride level [5]. Another prospective study for 4 years showed that weight gain more than 2 kg was related to increased risk of hypertension and hypercholesterolemia [31]. Most studies investigating the relationship between weight gain and metabolic health have been conducted in the general population, and a few studies have evaluated the relationship between weight change and MetS among immigrants [16]. As immigrants moved from their home country to the host country, they were likely to experience changes in dietary and lifestyle habits, resulting in a significant change in body weight [12,13]. A study of Asian immigrant women who migrated to Korea suggested that those who gained weight had an increased risk of MetS (defined as obesity (BMI ≥ 25 kg/m²) plus more than two of the following criteria: high blood pressure, high fasting glucose, hypertriglyceridemia, or low HDL cholesterol), although no quantitative analysis was conducted [16].
Previous studies have suggested that weight gain not only increased the risk of MetS, but also increased the risk of each component of MetS. In our study, the weight gain group had higher fasting glucose than the non-weight gain group. According to 15 year follow-up prospective study, subjects with more than 5 pounds weight gain had higher fasting glucose than the weight-maintaining group [32]. A randomized controlled trial for prevention of weight gain suggested that those who gained body weight 5% or more over 6 years had higher fasting glucose levels than those who maintained or lost body weight [33]. A retrospective study of the European population also supported the hypothesis that a change in BMI during early adulthood was associated with increased HbA1C [9]. However, another cross-sectional study showed that weight gain history was not associated with high fasting glucose [34].

Our study suggested significant positive relationship between weight gain and hypertriglyceridemia, which is consistent with previous studies [6,32,34]. A prospective study of the Caucasian and African American populations reported that normal-weight Caucasian men experienced a 16% increase in triglyceride level for each 9.1 kg of weight gain over 10 years [6]. Another cross-sectional study also suggested that weight gain during adulthood was related to high triglyceride level after controlling for current BMI [34].

There is controversy over the relationship between weight gain and blood pressure [6,34,35]. Prospective studies over 10 years showed that an average weight gain of 9.1 kg was related to 2.7–3.6 mmHg increase in blood pressure [6]. In contrast, another cross-sectional study found that weight gain was associated with risk of hypertension [34]. A longitudinal study over 5 years also observed no significant association between weight gain history and systolic or diastolic blood pressure in overweight African American population [35]. Our study found positive association between weight gain and high blood pressure among NKR in South Korea. Differences in length of follow-up, in levels on baseline BMI, or racial differences could influence the association between weight gain and blood pressure.

A significant association between weight gain and decrease in HDL cholesterol has been reported consistently in previous studies [6,32–34]. A prospective study suggested weight gain over 10 years was associated with decrease in HDL cholesterol from 0.09 mmol/L (Caucasian women) to 0.11 mmol/L (Caucasian men) [6]. Another prospective study over 15 years showed that weight gain was associated with decrease in HDL cholesterol level of 3.6 mg/dL (normal weight Caucasian population) to 3.8 mg/dL (overweight Caucasian population) [32]. A randomized controlled study for prevention of weight gain found that weight gain more than 5% had an adverse effect on HDL cholesterol [33]. Another cross-sectional study showed that weight gain in adulthood increased risk of low HDL cholesterol and high triglyceride [34]. However, we found no significant relationship between weight gain and low HDL cholesterol among NKR in South Korea. The differences between previous studies and our study might be due to racial differences, and further longitudinal studies are needed to clarify what has caused the differences of results between studies.

Although the weight gain group had higher BMI than the non-weight gain group on examination, the weight gain group had been thinner than the non-weight gain group in North Korea. One possible explanation for this is that NKR who experienced severe food shortage in North Korea might gain weight due to relative abundance of food in transit countries and South Korea [19,22]. NKR who experienced food shortage and severe malnutrition in the early life could benefit from increasing body muscle and fat mass after defection from North Korea. However, the thrifty phenotype hypothesis proposed poor nutrition in early life could produce altered glucose–insulin metabolism, which increases susceptibility to development of type 2 diabetes mellitus [36,37]. Furthermore, the weight gain group had a higher fat mass and percentage of body fat than the non-weight gain group on examination, suggesting that weight gain in the weight gain group after defection from North Korea may have led to increase in fat mass rather than muscle mass [38]. Studies have suggested that low muscle and high fat mass is related to insulin resistance.
and the risk of MetS [39–41]. Kim et al. demonstrated that higher muscle and lower fat mass were related to a lower insulin resistance and lower muscle and higher fat mass were associated with a higher insulin resistance and the risk of MetS (defined by modified NCEP-ATP III guideline) in the Korean population [39]. Srikanthan et al. also found that subjects in the highest quartile of skeletal muscle mass had a lower HOMA–IR value compared to those in the lowest quartile [40]. An increase in fat mass followed by weight gain over a relatively short time during resettlement in South Korea via transit countries could have increased insulin resistance and the risk of MetS among weight gain group in our study.

This study had several limitations. First, scientific sampling method was not applied in the NORNS study. Random sampling of NKRs in South Korea is difficult because only a few NKRs live in South Korea and they tend not to reveal their identities due to personal threats. However, the general demographics of participants in the NORNS study were similar to those of NKRs in South Korea who participated in a large-scale national survey in terms of gender ratio, age distribution, place of birth and stay duration, in transit countries [42]. Second, because of the cross-sectional design, no causal relationship between weight gain and risk of MetS could be determined. In addition, it was difficult to distinguish whether the difference in body composition between the weight gain group and the non-weight gain group was determined before migration or during migration and resettlement in South Korea. Third, this study used a self-reported questionnaire to measure weight change in NKRs subsequent to their defection from North Korea, although the validity and accuracy of self-reported measures of weight and weight change have been established in previous studies [43,44]. Fourth, NKRs have been in transit countries for different periods, which might have affected the risk of MetS. Fifth, we used BIA for body composition analysis. Although computed tomography (CT) or dual-energy X-ray absorptiometry (DXA) are the gold standard for evaluating distribution of body fat, BIA, which is highly correlated with CT and DXA, is the most frequently used method because of its simplicity, convenience, and low cost. [45,46]. Lastly, we did not implement a diet questionnaire in our survey, although there is a strong relationship between dietary patterns and MetS [47]. In the phase 2 study, we are collecting data by incorporating a diet questionnaire. In near future, our follow-up study will be able to elucidate the relationship between food consumption pattern and the risk of MetS in NKRs. We might better understand the effect of individual food and nutrients, such as whole-grain or soybean, on the risk of MetS [48,49]. Investigating relationship between changes in dietary pattern and MetS in NKRs will exhort to evaluate the changes in their microbiome among NKRs [50].

5. Conclusions

Our study suggested that the weight gain in NKRs during resettlement in South Korea increased the risk of MetS (defined as more than three out of five criteria: central obesity, impaired fasting glucose, elevated blood pressure, hypertriglyceridemia, and low HDL cholesterol) after adjusting for current BMI. An increase in body fat mass due to excess weight gain among NKRs might have an adverse effect on their metabolic health. Further prospective studies are needed to address the change in body weight of NKRs and its effect on the risk of cardiovascular disease and type 2 diabetes mellitus.

Supplementary Materials: The following are available online at https://www.mdpi.com/article/10.3390/ijerph18168479/s1, Table S1: Health questionnaire of NORNS study.

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References
1. Park, Y.W.; Zhu, S.; Palaniappan, L.; Heshka, S.; Carnethon, M.R.; Heymsfield, S.B. The metabolic syndrome: Prevalence and associated risk factor findings in the US population from the Third National Health and Nutrition Examination Survey, 1988–1994. *Arch. Intern. Med.* 2003, 163, 427–436. [CrossRef]
2. Mottillo, S.; Filion, K.B.; Genest, J.; Joseph, L.; Pilote, L.; Poirier, P.; Rinfret, S.; Schiffrin, E.L.; Eisenberg, M.J. The metabolic syndrome and cardiovascular risk: a systemic review and meta-analysis. *J. Am. Coll. Cardiol.* 2010, 56, 1113–1132. [CrossRef]
3. Wilson, P.W.; Dagostino, R.B.; Parise, H.; Sullivan, L.; Meigs, J.B. Metabolic syndrome as a precursor of cardiovascular disease and type 2 diabetes mellitus. *Circulation* 2005, 112, 3066–3072. [CrossRef] [PubMed]
4. Palaniappan, L.; Carnethon, M.R.; Wang, Y.; Hanley, A.J.; Fortmann, S.P.; Haffner, S.M.; Wagenknecht, L. Insulin Resistance Atherosclerosis Study. Predictors of the incident metabolic syndrome in adults: The Insulin Resistance Atherosclerosis Study. *Diabetes Care* 2004, 27, 788–793. [CrossRef]
5. Toga, S.; Fukkoshi, Y.; Akamatsu, R. Relationship between weight gain and metabolic syndrome in non-obese Japanese adults. *Diabetes Metab. Syndr.* 2016, 10, 63–67. [CrossRef]
6. Norman, J.E.; Bild, D.; Lewis, C.E.; Liu, K.; West, D.S. The impact of weight change on cardiovascular disease risk factors in young black and white adults: The CARDIA study. *Int. J. Obes. Relat. Metab. Disord.* 2003, 27, 369–376. [CrossRef]
7. Ouchi, N.; Parker, J.L.; Lugus, J.J.; Walsh, K. Adipokines in inflammation and metabolic disease. *Nat. Rev. Immunol.* 2011, 11, 85–97. [CrossRef]
8. Carnethon, M.R.; Loria, C.M.; Hill, J.O.; Sidney, S.; Savage, P.J.; Liu, K. Coronary Artery Risk Development in Young Adults study: Risk factors for the metabolic syndrome: The Coronary Artery Risk Development in Young adults (CARDIA) study, 1985–2001. *Diabetes Care* 2004, 27, 2707–2715. [CrossRef]
9. Montonen, J.; Boeing, H.; Schleicher, E.; Fritsche, A.; Pischon, T. Association of changes in body mass index during earlier adulthood and later adulthood with circulating obesity biomarker concentrations in middle-aged men and women. *Diabetologia* 2011, 54, 1676–1683. [CrossRef]
10. Hashimoto, Y.; Hamaguchi, M.; Fukuda, T.; Obora, A.; Kojima, T.; Fukui, M. Weight gain since age of 20 as risk of metabolic syndrome even in non-overweight individuals. *Endocrine* 2017, 58, 253–261. [CrossRef]
11. Suzuki, A.; Akamatsu, R. Long-term weight gain is related to risk of metabolic syndrome even in non-obese individuals. *Endocrine* 2017, 58, 253–261. [CrossRef]
12. Gadd, M.; Sundquist, J.; Johansson, S.E.; Wandell, P. Do immigrants have an increased prevalence of unhealthy behaviours and risk factors for coronary heart disease? *Eur. J. Cardiovasc. Prev. Rehabil.* 2005, 12, 535–541. [CrossRef]
13. Tseng, M.; Wright, D.J.; Fang, C.Y. Acculturation and dietary change among Chinese immigrant women in United States. *J. Immigr. Minor. Health* 2015, 17, 400–407. [CrossRef]
14. Sattar, N.; Gill, J.M.R. Type 2 diabetes in migrant South Asians: Mechanisms, mitigation and management. *Lancet Diabetes Endocrinol.* 2015, 3, 1004–1016. [CrossRef]
15. Silbiger, J.J.; Stein, R.; Roy, M.; Nair, M.K.; Cohen, P.; Shaffer, J.; Pinkhasov, A.; Kamran, M. Coronary artery disease in South Asian immigrants living in New York City: Angiographic findings and risk factors burdens. *Etnh. Dis.* 2013, 23, 292–295.
16. Yang, S.J.; Chee, Y.K.; Kim, J.A.; An, J. Metabolic syndrome and its related factors among Asian immigrant women in Korea. *Nurs. Health Sci. Health* 2014, 16, 373–380. [CrossRef] [PubMed]
17. STEPWise Approach to Chronic Disease Risk Factor Surveillance; World Health Organization: Geneva, Switzerland, 2005. Available online: https://www.who.int/teams/noncommunicable-diseases/surveillance/data/democratic-people-s-republic-of-korea (accessed on 30 March 2021).
18. Kang, Y.H. Two Koreas, war and health. *Int. J. Epidemiol.* 2013, 42, 925–929. [CrossRef] [PubMed]
19. McCurry, J. No end in sight for North Korea’s malnutrition crisis. *Lancet* 2012, 379, 602. [CrossRef]
20. Lee, Y.; Lee, M.K.; Chun, K.H.; Lee, Y.K.; Yoon, S.J. Trauma experience of North Korean refugees in China. *Am. J. Prev. Med.* 2001, 20, 225–229. [CrossRef]

21. Current Status of North Korean Refugees’ Resettlement; Ministry of Unification: Seoul, Korea, 2021. Available online: http://unikorea.go.kr/unikorea/business/NKDefectorsPolicy/status/lately (accessed on 30 March 2021).

22. Kim, Y.J.; Kim, S.G.; Lee, Y.H. Prevalence of general and central obesity and associated factors among North Korean refugees in South Korea by duration after defection from North Korea: A cross-sectional study. *Int. J. Environ. Res. Public Health* 2018, 15, 811. [CrossRef]

23. Kim, Y.J.; Lee, Y.H.; Lee, Y.J.; Kim, K.J.; An, J.H.; Kim, N.H.; Kim, H.Y.; Choi, D.S.; Kim, S.G. Prevalence of metabolic syndrome and its related factors among North Korean refugees in South Korea: A cross-sectional study. *BMJ Open* 2016, 6, e010849. [CrossRef] [PubMed]

24. Lee, Y.H.; Lee, W.J.; Kim, Y.J.; Cho, M.J.; Kim, J.H.; Lee, Y.J.; Kim, H.Y.; Choi, D.S.; Kim, S.G.; Robinson, C. North Korean refugee health in South Korea (NORNS) study: Study design and methods. *BMC Public Health* 2012, 12, 172. [CrossRef]

25. Ma, W.-Y.; Yang, C.-Y.; Shih, S.-R.; Hsieh, H.-J.; Hung, C.S.; Chiu, F.-C.; Lin, M.-S.; Liu, P.-H.; Hua, C.-H.; Hsein, Y.-C.; et al. Measurement of waist circumference: Midabdominal or iliac crest? *Diabetes Care* 2013, 36, 1660–1666. [CrossRef] [PubMed]

26. Kobalava, Z.D.; Kotovskaya, Y.V.; Babaeva, L.A.; Moiseev, V.S. Validation of TM-2655 oscillometric device for blood pressure measurement. *Blood Press. Monit.* 2006, 11, 87–90. [CrossRef]

27. Matthews, D.R.; Hosker, J.P.; Rudenski, A.S.; Naylor, B.A.; Treacher, D.F.; Turner, R.C. Homeostasis model assessment: Insulin resistance and beta-cell function from fasting plasma glucose and insulin concentrations in man. *Diabetologia* 1985, 28, 412–419. [CrossRef]

28. Grundy, S.M.; Cleeman, J.I.; Daniels, S.R.; Donato, K.A.; Eckel, R.H.; Franklin, B.A.; Gordon, D.J.; Krauss, R.M.; Savage, P.J.; Smith, S.C.; et al. American Heart Association; National Heart, Lung, and Blood Institute. Diagnosis and management of the metabolic syndrome: An American Heart Association/National Heart, Lung, and Blood Institute Scientific Statement. *Circulation* 2005, 112, 2735–2752. [CrossRef]

29. Lewis, C.E.; Jacobs, D.R.; McCreath, H.; Kiefte, C.I.; Schreiner, P.J.; Smith, D.E.; Williams, O.D. Weight gain continues in the 1990s: 10-year trends in weight and overweight from the CARDIA study. Coronary Artery Risk Development in Young Adults. *Am. J. Epidemiol.* 2000, 151, 1172–1181. [CrossRef]

30. Willett, W.C.; Manson, J.E.; Stampfer, M.J.; Colditz, G.A.; Rosner, B.; Speizer, F.E.; Hennekens, C.H. Weight, weight change, and coronary heart disease in women. Risk within the ‘normal’ weight range. *JAMA* 1995, 273, 461–465. [CrossRef] [PubMed]

31. Ishikawa-Takata, K.; Ohta, T.; Moritaki, K.; Gotou, T.; Inoue, S. Obesity, weight change and risks for hypertension, diabetes and hypercholesterolemia in Japanese men. *Eur. J. Clin. Nutr.* 2002, 56, 601–607. [CrossRef]

32. Truesdale, K.P.; Stevens, J.; Lewis, C.E.; Schreiner, P.J.; Smith, C.M.; Cai, J. Changes in risk factors for cardiovascular disease by baseline weight status in young adults who maintain or gain weight over 15 years: The CARDIA study. *Int. J. Obes.* 2006, 30, 1397–1407. [CrossRef]

33. Wing, R.R.; Espeland, M.A.; Tate, D.F.; Perdue, L.H.; Bahnsen, J.; Polzien, K.; Robichaud, E.F.; LaRose, J.G.; Gorin, A.A.; Lewis, C.E.; et al. Study of Novel Approaches to Weight Gain Prevention Research Group. Changes in cardiovascular risk factors over 6 years in young adults in a randomized trial of weight gain prevention. *Obesity* 2020, 28, 2323–2330. [CrossRef]

34. Alley, D.E.; Chang, V.W. Metabolic syndrome and weight gain in adulthood. *J. Gerontol. A Biol. Sci. Med. Sci.* 2010, 65, 111–117. [CrossRef] [PubMed]

35. Curtis, A.B.; Strogatz, D.S.; James, S.A.; Raghunathan, T.E. The contribution of baseline weight and weight gain to blood pressure change in African Americans: The Pitt County Study. *Ann. Epidemiol.* 1995, 8, 497–503. [CrossRef] [PubMed]

36. Hales, C.N.; Barker, D.J. Type 2 (non-insulin-dependent) diabetes mellitus: The thrifty phenotype hypothesis. *Diabetologia* 1992, 35, 595–601. [CrossRef]

37. Hales, C.N.; Barker, D.J. The thrifty phenotype hypothesis. *Br. Med. Bull.* 2001, 60, 5–20. [CrossRef]

38. Cho, S.W.; Lee, S.H.; Koh, E.S.; Kim, S.E.; Kim, S.J. Characteristics of body composition and muscle strength of North Korean refugees during South Korean Stay. *Endocrinol. Metab.* 2015, 30, 551–556. [CrossRef] [PubMed]

39. Kim, K.; Park, S.M. Association of muscle mass and fat mass with insulin resistance and the prevalence of metabolic syndrome in Korean adults: A cross-sectional study. *Sci. Rep.* 2018, 8, 2703. [CrossRef] [PubMed]

40. Srikanthan, P.; Karlamangla, A.S. Relative muscle mass is inversely associated with insulin resistance and prediabetes. Findings from the third National Health and Nutrition Examination Survey. *J. Clin. Endocrinol. Metab.* 2011, 96, 2898–2903. [CrossRef] [PubMed]

41. Kim, G.; Lee, S.; Jun, J.E.; Lee, Y.; Ahn, J.; Bae, J.C.; Jin, S.; Hur, K.Y.; Lee, J.H.; Lee, M.; et al. Increase in relative skeletal muscle mass over time and its inverse association with metabolic syndrome development: A 7-year retrospective cohort study. *Cardiovasc. Diabetol.* 2018, 17, 23. [CrossRef]

42. Current Stats Survey on North Korean Refugees Living in South Korea (Korean); North Korean Refugee Foundation: Seoul, Korea, 2012.
45. Xu, L.; Cheng, X.; Wang, J.; Cao, Q.; Sato, T.; Wang, M.; Zhao, X.; Liang, W. Comparisons of body-composition prediction accuracy: A study of 2 bioelectric impedance consumer devices in healthy Chinese persons using DXA and MRI as criteria methods. *J. Clin. Densitom.* 2011, 14, 458–464. [CrossRef] [PubMed]

46. Thomson, R.; Brinkworth, G.D.; Buckley, J.D.; Noakes, M.; Clifton, P.M. Good agreement between bioelectrical impedance and dual-energy X-ray absorptiometry for estimating changes in body composition during weight loss in overweight young women. *Clin. Nutr.* 2007, 26, 771–777. [CrossRef] [PubMed]

47. Kastorini, C.M.; Milionis, H.J.; Esposito, K.; Giugliano, D.; Goudevenos, J.A.; Panagiotakos, D.B. The effect of Mediterranean diet on metabolic syndrome and its components: A meta-analysis of 50 studies and 534,906 individuals. *J. Am. Coll. Cardiol.* 2011, 57, 1299–1313. [CrossRef]

48. Sahyoun, N.R.; Jacques, P.F.; Zhang, X.L.; Juan, W.; McKeown, N.M. Whole-grain intake is inversely associated with the metabolic syndrome and mortality in older adults. *Am. J. Clin. Nutr.* 2006, 83, 124–131. [CrossRef]

49. Goodman-Gruen, D.; Kritz-Silverstein, D. Usual dietary isoflavone intake is associated with cardiovascular disease risk factors in postmenopausal women. *J. Nutr.* 2001, 131, 1202–1206. [CrossRef]

50. Dabke, K.; Hendrick, G.; Devkota, S. The gut microbiome and metabolic syndrome. *J. Clin. Investig.* 2019, 129, 4050–4057. [CrossRef]