## RESEARCH

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# The impact of weight change on suicide mortality: a nationwide population-based cohort study of 2 million Koreans

Kyuho Kim<sup>1</sup>, Jin-Hyung Jung<sup>2</sup>, Yoo Hyun Um<sup>3</sup>, Yu-Bae Ahn<sup>1</sup>, Seung-Hyun Ko<sup>1</sup>, Kyungdo Han<sup>4\*†</sup> and Jae-Seung Yun<sup>1\*†</sup>

## Abstract

**Background** Previous studies have shown that weight change has a reverse J-shape association with all-cause mortality. However, its association with suicide mortality remains undetermined. In this study, we investigated the association between weight change and suicide mortality using a large-scale, population-based cohort from the Korean National Health Insurance Service database.

**Methods** A total of 2,103,525 subjects aged  $\geq$  20 years who underwent a general health screening program twice in the 2-year interval between 2007 and 2009 were included. Subjects were categorized into five groups according to the percent weight change during this period: severe weight loss (< - 15.0%), moderate weight loss (- 15.0%), weight stable (- 5.0 to < 5.0%), moderate weight gain (5.0 to < 15.0%), and severe weight gain ( $\geq$  15.0%).

**Results** During a median follow-up of 11.3 years, 6,179 cases (0.3%) of suicide mortality occurred. Weight change was associated with increased suicide mortality in a reverse J-shaped curve, even after adjustment for covariates. In particular, those with severe weight loss or gain showed 1.8-fold or 1.6-fold increased risk of suicide mortality, respectively. This reverse J-shaped association was consistently observed in subgroup analyses considering age, sex, depression, cancer, and BMI category.

**Conclusions** Moderate to severe weight change within a 2-year interval is associated with increased risk of suicide mortality. To better understand the mechanisms through which weight change affects suicide mortality, studies incorporating information on weight change intentions, medications, weight change-related medical conditions are needed.

Keywords Mortality, Suicide, Weight change, Weight gain, Weight loss

 $^{\dagger}\mbox{Kyungdo}$  Han and Jae-Seung Yun have contributed equally to this study as co-corresponding authors.

<sup>3</sup> Department of Psychiatry, St. Vincent's Hospital, College of Medicine, The Catholic University of Korea, Seoul, Republic of Korea

<sup>4</sup> Department of Statistics and Actuarial Science, Soongsil University, 369 Sangdo-ro, Dongjak-gu, Seoul 06987, Republic of Korea

\*Correspondence: Kyungdo Han hkd917@naver.com Jae-Seung Yun dryun@catholic.ac.kr <sup>1</sup> Division of Endocrinology and Metabolism, Department of Internal Medicine, St. Vincent's Hospital, College of Medicine, The Catholic University of Korea, 222 Banpo-daero, Seocho-gu, Seoul 06591, Republic

University of Korea, 222 of Korea

<sup>2</sup> Samsung Biomedical Research Institute, Sungkyunkwan University School of Medicine, Suwon, Republic of Korea



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## Introduction

Overweight or obesity is a risk factor for cardiovascular disease (CVD) and metabolic diseases such as type 2 diabetes, hypertension, and dyslipidemia [1, 2]. Body mass index (BMI) is a popular indicator of an individuals' fatness [3], and several studies have consistently shown that both high BMI (overweight or obesity) and low BMI (underweight) are associated with increased mortality, following J-shaped or U-shaped curves [4–6].

Whether weight loss or weight gain can influence mortality is an intriguing question; several studies have indeed demonstrated that both weight loss [7, 8] and weight gain [9, 10] are associated with increased mortality. In addition, a nationwide population-based study including approximately 10 million Korean adults demonstrated that weight loss or gain > 5.0% within a 4-year interval is associated with increased risk of all-cause mortality [11]. However, the association between weight change and cause-specific mortality has not been studied well.

According to the World Health Organization (WHO), suicide accounted for 1.3% of all deaths in 2019 [12]. In United States, it was the 10th leading cause of death overall, the 2nd in people aged 10-34 years, and the 5th in people aged 35-54 years [13]. In South Korea, it is the 5th leading cause of death overall, the 1st in people aged 10–39, and 2nd in people aged 40-59 years [14]. A previous prospective cohort study including 18,784 men aged 40-69 years in the United Kingdom showed unexplained weight loss to be associated with an increased risk of suicide mortality [15]. However, the study was limited in that only men were included, and information was lacking about the amount of weight loss and diagnoses of psychiatric disorders such as depression. In addition, research is scarce concerning how much weight change increases suicide mortality among subjects of differing BMI categories. Considering that the European Medicines Agency has started to review data on the risk of suicidal thoughts with anti-obesity medications such as semaglutide and liraglutide [16], there is a need to examine whether weight change is associated with suicide mortality according to BMI category.

Therefore, we aimed to investigate the association between weight change and suicide mortality using a large-scale, population-based cohort derived from the Korean National Health Insurance Service (NHIS) general health screening program. In addition, we performed subgroup analyses according to age, sex, diabetes, depression, cancer, and BMI category.

## Methods

## Study population

The NHIS is a mandatory national health insurance managed by the Korean government, providing health insurance to approximately 97.0% of the Korean population. The NHIS database includes data on demographics, diagnoses, prescriptions, medical procedures, inpatient and outpatient service records, and death records. The NHIS offers a general health screening program biannually for all insured Koreans, with the exception of non-office workers who receive the general health screening program annually [17]. The proportion of participants in the general health screening program relative to the eligible population has been approximately 75% over the past ten years [18]. The general health screening program includes tests for cardiovascular and cerebrovascular diseases, including hypertension, diabetes mellitus, dyslipidemia, and obesity, and for other diseases such as liver disease, psychiatric disorders, and geriatric diseases [19].

We identified 4,234,415 subjects aged  $\geq$  20 years who underwent the general health screening program through the NHIS from January 1, 2009 to December 31, 2009 (index date). Among these, we included subjects who had also participated in the general health screening program 2 years before (in 2007) (N=2,380,806). We excluded subjects with missing data for one or more variables in the general health screening program at the index date (N=272,763). In addition, we excluded subjects who died within 1 year after cohort entry (N=4,518). Ultimately, a total of 2,103,525 subjects were included and followed until the occurrence of suicide death, or December 31, 2021 (Fig. 1).

This study was approved by the Institutional Review Board of The Catholic University of Korea, St. Vincent's Hospital (VC23ZISE0218), and informed consent was waived. The study was conducted in accordance with the principles of the Declaration of Helsinki.

## Definition of weight change

Weight change was calculated as percent change between 2007 and 2009:  $(100 \times [\text{weight at } 2009-\text{weight at } 2007]/\text{weight at } 2007]$ . We categorized subjects into five groups based on weight change, with reference to previous studies [11, 20, 21]: severe weight loss (weight change < - 15.0%), moderate weight loss (weight change - 15.0 to < - 5.0%), weight stable (weight change - 5.0 to < 5.0%), moderate weight gain (weight change 5.0 to < 15.0%), and severe weight gain (weight change  $\geq 15.0\%$ ).



Fig. 1 A flowchart of the study population enrollment. NHIS denotes National Health Insurance Service

## Definition of variables and end point

Demographic data, anthropometric measurements, laboratory results, and comorbidities were obtained during the general health screening program in 2009 (index date). Healthcare professionals measured anthropometric indices (BMI, waist circumference, and blood pressure [BP]) during the general health screening program. Blood pressure was measured using a standard sphygmomanometer after at least 5 min of rest. Blood samples were collected after overnight fasting, and serum glucose and total cholesterol were measured. Information about smoking status, alcohol consumption, and exercise were obtained using selfreported questionnaires. Smoking status was classified as nonsmoker, ex-smoker, or current smoker. Alcohol drinker was defined when subjects had alcohol intake >0 g per day. Regular exercise was defined as high-intensity activities  $\geq 3$  times a week or moderate intensity activities  $\geq 5$  times a week. Income was classified into quartiles (Q): Q1 (low income), Q2, Q3, and Q4. Obesity was defined as a BMI of  $\geq 25$  kg/m<sup>2</sup> according to the guidelines of the Korean Society for the Study of Obesity [22]. Diabetes was defined as at least one claim per year for the prescription of antidiabetic medications under the International Statistical Classification of Diseases, 10th revision (ICD-10) codes E11-E14, or fasting glucose level  $\geq$  126 mg/dL at the time of health screening. Hypertension was defined as systolic BP  $\geq$  140 mmHg, diastolic BP  $\geq$  90 mmHg, the ICD-10 codes I10-13 or I15, or prescription of antihypertensive medications. Dyslipidemia was defined as a total cholesterol level  $\geq$  240 mg/dL, the ICD-10 code E78, or prescription of lipid-lowering medications. Chronic kidney disease (CKD) was defined based on the estimated glomerular filtration rate (eGFR) calculated by the Modification of Diet in Renal Disease formula, with eGFR < 60 mL/min/1.73 m<sup>2</sup> considered CKD. Depression was defined as the ICD-10 code F32 or F33, as previously described [23-25]. Eating disorder was defined as the ICD-10 code F50. Substance use disorder was defined as the ICD-10 code F10-F19. The NHIS established a special co-payment service to enhance health coverage and relieve the financial burden of patients with cancer. Because enrollment in this co-payment service is indicated by a special reimbursement code for cancer (V193) and requires a medical certificate from a physician, the cancer diagnoses are considered to be sufficiently reliable. Therefore, cancer was defined using the ICD-10 'C' code and the cancer reimbursement code (V193), as previously described [26, 27].

The endpoint was suicide mortality during the followup period. Suicide deaths were identified by ICD-10 codes X60–X84, as recorded in the government mortality database (Statistics Korea), as previously described [28]. Follow-up duration was defined as the interval from the index date to the date of suicide mortality, or until December 2021, whichever came first.

## Statistical analysis

Data were expressed as the mean±standard deviation, median (interquartile range), or number with percentage (%). Baseline characteristics were compared between groups using ANOVA for continuous variables and the chi-square test for categorical variables. The incidence rate (IR) of suicide mortality was calculated by dividing the number of events per 1,000 person-years. Hazard ratios (HRs) and 95% confidence intervals (CIs) for suicide mortality according to the 5 weight change categories were calculated using multivariable Cox proportional hazard regression models.

Model 1 was non-adjusted. Model 2 was adjusted for age, sex, income (Q1), smoking, alcohol drinking, and regular exercise. Model 3 was adjusted for model 2 plus diabetes, hypertension, dyslipidemia, CKD, and depression. Model 4 was adjusted for model 3 plus weight at the 2009 screening [29]. Model 5 was adjusted for model 4 plus eating disorder and substance use disorder. Subgroup analyses were conducted based on age, sex, diabetes, depression, and cancer, after adjusting for age, sex, income (Q1), smoking, alcohol consumption, regular exercise, diabetes, hypertension, dyslipidemia, chronic kidney disease, depression, and weight at the 2009 screening. Given that waist circumference was positively associated with mortality across all BMI categories examined [30], waist circumference was adjusted instead of weight at the 2009 screening in the BMI subgroup analysis.

All analyses were performed using SAS software version 9.3 (SAS Institute, Cary, NC, USA). A two-sided P < 0.05 was considered statistically significant without adjustment for multiple comparisons.

## Results

## **Baseline characteristics**

Subjects were  $48.0 \pm 13.6$  years old, with body weight of  $64.4 \pm 11.5$  kg and BMI of  $23.8 \pm 3.1$  kg/m<sup>2</sup>. Obesity was present in 695,159 (33.1%) subjects, diabetes in 180,616 (8.6%), hypertension in 545,535 (25.9%), dyslipidemia in 375,378 (17.9%), CKD in 156,910 (7.5%), and depression in 62,978 (3.0%). Subjects in the weight loss groups were older, had higher fasting glucose levels, lower total cholesterol levels, and were less likely to be smokers or alcohol drinkers. Additionally, they had a higher prevalence of diabetes, hypertension, CKD, and cancer compared to those in the weight stable group. Subjects in the weight gain groups were younger, had lower fasting glucose levels, higher total cholesterol levels, and were less likely to be regular exercisers. Additionally, they had a lower prevalence of diabetes and hypertension compared to those in the weight stable group. Compared to those in the weight stable group, subjects in the weight gain or weight loss groups were less likely to be male and had a higher prevalence of low income (Q1) status and depression (Table 1).

## Suicide mortality according to weight change

Over a median follow-up of 11.3 years (interquartile range, 11.1-11.6), 6,179 cases of suicide mortality occurred. The weight loss group had a higher incidence of suicide mortality than the weight stable or weight gain groups (Additional file 1: Fig. S1). The IR of suicide death was 0.58, 0.35, 0.25, 0.26, and 0.31 per 1,000 person-years for the severe weight loss, moderate weight loss, weight stable, moderate weight gain, and severe weight gain groups, respectively. In the non-adjusted model, severe or moderate weight loss associated with increased risk of suicide mortality. After multivariable adjustment, suicide mortality showed a reverse J-shaped curve according to weight change. Both weight loss (HR, 1.84; 95% CI 1.44-2.36 for the severe weight loss group, and HR, 1.23; 95% CI 1.14-1.32 for the moderate weight loss group) and weight gain (HR, 1.62; 95% CI 1.26-2.07 for the severe weight gain group, and HR, 1.23; 95% CI 1.14-1.33 for the moderate weight gain group) were significantly associated with increased risk of suicide mortality compared with the weight stable group (Table 2 and Fig. 2).

# Subgroup analyses according to age, sex, diabetes, depression, cancer, and BMI category

The relationship between weight change and risk of suicide mortality did not demonstrate significant interactions with age, sex, depression, cancer, or BMI category, instead showing a consistent trend across these factors. However, a significant interaction between diabetes presence and suicide risk was found (P = 0.03 for interaction), with no significant increase in risk for severe weight loss among patients with diabetes. Conversely, a 1.9-fold increase in the risk of suicide mortality was observed in those with diabetes and severe weight gain (Additional file 1: Table S1 and Figs. S2, S3, S4).

## Discussion

In this study, we investigated the association between body weight changes and suicide mortality. The main findings are summarized as follows. First, approximately 25.6% and 1.4% of this large nation-wide population-based cohort experienced moderate or severe weight changes respectively in the 2-year follow-up period. Second, a reverse J-shaped association between weight change and suicide mortality was observed; especially, those with severe weight loss or weight gain showed 1.8-fold or 1.6-fold increased risk of suicide mortality, respectively. Third, this trend of U-shaped or J-shaped association was consistently observed in subgroup analyses according to sex, age, depression, cancer, and BMI category.

#### Total Severe weight Moderate weight Weight stable Moderate Severe weight P for trend loss (weight loss (weight (weight change weight gain gain (weight change < - 15.0%) change – 15.0 - 5.0 to < 5.0%) (weight change change $\geq$ 15.0%) to <- 5.0%) 5.0% to < 15.0%) (N = 2,103,525) (N = 10,713) (N=219,740) (N=1,565,679) (N = 288,375) (N = 19,018) Age (years) $48.0 \pm 13.6$ $51.6 \pm 17.5$ $50.1 \pm 14.9$ $48.3 \pm 13.2$ $44.7 \pm 13.8$ $43.8 \pm 16.2$ < 0.001 Age group—no. < 0.001 (%) 20-39 640,919 (30.5) 3,349 (31.3) 60,294 (27.4) 446,106 (28.5) 121,446 (42.1) 9,724 (51.1) 40-64 914,676 (58.4) 1,178,114 (56.0) 4,306 (40.2) 116,316 (52.9) 136,355 (47.3) 6,461 (34.0) ≥65 284,492 (13.5) 3,058 (28.5) 43,130 (19.6) 204,897 (13.1) 30,574 (10.6) 2,833 (14.9) Men—no (%) 1,243,764 (59.1) 4.632 (43.2) 114,786 (52,2) 946.221 (60.4) 168,786 (58,5) 9.339 (49.1) < 0.001 Height (cm) $164.3 \pm 9.2$ $160.5 \pm 9.7$ $162.7\pm9.4$ $164.5 \pm 9.1$ $165.0 \pm 9.4$ $163.8 \pm 9.5$ < 0.001 Weight (kg) 64.4±11.5 $54.6 \pm 10.8$ $60.0 \pm 10.8$ 64.6±11.3 $66.9 \pm 12.0$ 69.7±13.1 < 0.001 $21.1 \pm 3.1$ $23.8 \pm 3.1$ $25.9 \pm 3.9$ BMI (kg/m<sup>2</sup>) $23.8 \pm 3.1$ $22.6 \pm 3.0$ $24.5 \pm 3.2$ < 0.001 BMI (kg/m<sup>2</sup>) $23.7 \pm 3.1$ $26.2 \pm 4.1$ $24.4 \pm 3.2$ $23.8 \pm 3.1$ $22.8 \pm 3.0$ $21.4 \pm 3.0$ < 0.001 in 2007 Waist circumfer- 80.7 ± 8.9 $77.2 \pm 9.6$ $78.2 \pm 9.7$ $80.8\pm8.8$ $81.8 \pm 8.9$ $83.6 \pm 9.8$ < 0.001 ence (cm) Fasting glucose 97.2±22.8 $100.5 \pm 37.8$ $98.7 \pm 29.7$ $97.2 \pm 22.1$ $95.7 \pm 19.6$ $95.7 \pm 22.7$ < 0.001 (mg/dL)Total choles- $195.9 \pm 36.4$ $187.9 \pm 37.7$ $191.6 \pm 36.9$ $196.2 \pm 36.2$ 197.6 + 36.7202.1 + 40.5< 0.001 terol (mg/dL) Systolic BP $122.7 \pm 14.7$ $121.8 \pm 15.9$ $121.8 \pm 15.1$ $122.8 \pm 14.6$ $122.8 \pm 14.6$ $122.7 \pm 15.2$ < 0.001 (mmHg) Diastolic BP $76.6 \pm 9.9$ $75.8 \pm 9.9$ $76.7 \pm 9.8$ $76.7 \pm 9.9$ < 0.001 $75.5 \pm 10.2$ $76.6 \pm 10.1$ (mmHg) I ow-income 329,169 (15.7) 2,033 (19.0) 39,615 (18.0) 239,415 (15.3) 44,879 (15.6) 3,227 (17.0) < 0.001 status—no. (%) Current 551,756 (26.2) 2,109 (19.7) 53,945 (24.6) 412,633 (26.4) 78,628 (27.3) 4,441 (23.4) < 0.001 smoker—no. (%) Alcohol < 0.001 1,048,129 (49.8) 3,836 (35.8) 96,321 (43.8) 790,636 (50.5) 149,068 (51.7) 8.268 (43.5) drinker—no. (%) Regular exer-402,197 (19.1) 2,109 (19.7) 45,412 (20.7) 305,685 (19.5) 46,396 (16.1) 2,595 (13.6) < 0.001 cise-no. (%) Obesity-no. 695,159 (33.1) 1,100 (10.3) 43,238 (19.7) 522,313 (33.4) 118,112 (41.0) 10.396 (54.7) < 0.001 (%) Diabetes—no. 180,616 (8.6) 132,886 (8.5) 1,589 (8.4) < 0.001 1,501 (14.0) 25,312 (11.5) 19,328 (6.7) (%) Hyperten-545,535 (25.9) 3.376 (31.5) 60,850 (27.7) 410,051 (26.2) 66,662 (23.1) 4,596 (24.2) < 0.001 sion-no. (%) Dyslipidemia— 375,378 (17.9) 1,781 (16.6) 38,240 (17.4) 280,856 (17.9) 50,419 (17.5) 4,082 (21.5) < 0.001 no. (%) CKD—no. (%) 156,910 (7.5) 1,078 (10.1) 17,373 (7.9) 116,728 (7.5) 20,240 (7.0) 1,491 (7.8) < 0.001 Depression— 62,978 (3.0) 602 (5.6) 9,473 (4.3) 43,184 (2.8) 8,974 (3.1) 745 (3.9) < 0.001 no. (%) Eating disor-700 (0.0) 7 (0.1) 123 (0.1) 459 (0.0) 98 (0.0) 13 (0.1) < 0.001 der-no. (%) Substance use 4,470 (0.2) 36 (0.3) 667 (0.3) 3,017 (0.2) 680 (0.2) 70 (0.4) < 0.001 disorder—no. (%) 530 (5.0) 4,297 (2.0) 3,153 (1.1) 237 (1.3) < 0.001 Cancer—no. 26,584 (1.3) 18,367 (1,2) (%)

## Table 1 Baseline characteristics of the study population in 2009

Data are shown as mean ± standard deviation, median (25–75th percentile), or number (%). BMI denotes body mass index, BP blood pressure, and CKD chronic kidney disease

Weight change	Number	Event	IR/1,000 PY	Model 1	Model 2	Model 3	Model 4	Model 5
				HR (95% CI)				
<-15.0%	10,713	64	0.58	2.31 (1.80–2.95)	2.34 (0.83–2.99)	2.25 (1.76–2.88)	1.84 (1.44–2.36)	1.83 (1.43–2.35)
− 15.0 to < − 5.0%	219,740	832	0.35	1.38 (1.29–1.49)	1.36 (1.26–1.46)	1.33 (1.23–1.43)	1.23 (1.14–1.32)	1.22 (1.14–1.32)
- 5.0 to < 5.0%	1,565,67	9 4,376	0.25	1 (Ref.)				
5.0 to < 15.0%	288,375	843	0.26	1.05 (0.97–1.13)	1.21 (1.13–1.30)	1.19 (1.11–1.28)	1.23 (1.14–1.33)	1.23 (1.14–1.32)
≥15.0%	19,018	64	0.31	1.23 (0.96–1.57)	1.53 (1.20–1.96)	1.48 (1.16–1.90)	1.62 (1.26–2.07)	1.59 (1.25–2.04)

Table 2 Association between weight change and suicide mortality

Model 1: non-adjusted. Model 2: model 1 plus adjustment for age, sex, income (Q1), smoking, alcohol drinking, and regular exercise. Model 3: model 2 plus adjustment for diabetes, hypertension, dyslipidemia, chronic kidney disease, and depression. Model 4: model 3 plus adjustment for weight at the 2009 screening. Model 5: model 4 plus adjustment for eating disorder and substance use disorder

CI denotes confidence interval, HR hazard ratio, IR incidence rate, and PY person-years



Weight change category over 2 years

Fig. 2 Risk of suicide mortality. Hazard ratios (HRs) with 95% confidence intervals (Cls) are presented as dot and whisker plots after adjusting for covariates: age, sex, income (Q1), smoking, alcohol drinking, regular exercise, diabetes, hypertension, dyslipidemia, chronic kidney disease, depression, weight at the 2009 screening, eating disorder, and substance use disorder. Severe weight loss group (weight change < -15.0%), moderate weight loss group (weight change – 15.0%), weight stable group (weight change – 5.0%), moderate weight change 5.0 to < 15.0%), and severe weight gain (weight change ≥ 15.0%)

Several previous studies have shown that weight change, whether loss or gain, is associated with increased mortality [11, 31, 32]. However, only a few studies have considered the association between weight change and suicide mortality. Considering that only 24.0% of those who die by suicide had psychiatric diagnoses in the 4-week period prior to death [33], it is helpful to identify those at high risk of suicide based on physiologic indicators along with psychiatric markers. The reverse J-shaped association between weight change and suicide mortality observed in this study, which was independent of depression, suggests that weight change can be used as physiologic indicator for suicide mortality. From another perspective, considering the rise in popularity of semaglutide [34] and bariatric surgery [35] and the recent release of tirzepatide [36], all of which provide for weight loss of  $\geq 15.0\%$  in subjects with obesity, it is meaningful to examine the association between weight change and suicide mortality. Due to the large number of subjects and long-term follow-up period, the Korean nationwide database provides a good opportunity to investigate the association between weight change and suicide mortality.

Our observation of increased risk of suicide mortality in association with weight loss is in line with previous findings [15, 37]. A prospective cohort study in the United Kingdom showed unexplained weight loss to be associated with increased risk of suicide mortality [15]. However, that study included only men aged 40-69, and it was limited by not considering depression. Another study in Japan, which analyzed a cohort of 41,746 adults and identified 80 cases of suicide events, revealed weight loss since the age of 20 to be significantly associated with an elevated risk of suicide mortality [38]. Several hypotheses can be suggested as potential reasons for suicide resulting from weight loss. Mental health issues such as appetite suppression due to depression, significant psychological stress, and feelings of lethargy can lead to sudden weight loss, potentially triggering thoughts and behaviors related to suicide [37]. Severe systemic illnesses such as cancer, infectious diseases, and gastrointestinal disorders can also be associated with weight loss. A previous study using data from the National Health and Nutrition Examination Survey demonstrated that subjects losing at least 10% of weight within one year have likely been diagnosed with medical illnesses such as arthritis, diabetes, cancer, cardiovascular disease, liver conditions, and respiratory diseases [39]. Dietary restrictions can also lead to severe weight loss. Our study

utilized the NHIS general health screening cohort, which included relatively healthy individuals who underwent consecutive biennial national health examinations. In our findings, rapid weight loss was significantly associated with increased risk of suicide even in individuals without a history of cancer, major metabolic diseases such as diabetes, or depression. In addition, subgroup analyses demonstrated that regardless of baseline BMI category, i.e., normal weight, overweight, or obesity, moderate or severe weight change is associated increased risk of suicide mortality. Conversely, among young adults aged 20-39, those with diabetes, depression, cancer, or obesity class II showed no statistically significant increase in suicide risk associated with severe weight loss. The interpretation of these results may consider factors such as a low occurrence of suicide deaths in these groups, deaths due to factors other than suicide, and potential health improvements resulting from weight loss.

We also found increased risk of suicide mortality to be associated with weight gain in this study. One possible explanation is that weight control failure (weight gain despite weight loss efforts) could increase risk of suicidal ideation [40]. In addition, considering that psychiatric disorders such as schizophrenia [41] and bipolar disorder [42] are risk factors of suicide, and medications for these disorders increases body weight [43, 44], increased suicide mortality associated with weight gain might be attributed to psychiatric disorders. Whereas subjects without diabetes showed a reverse J-shaped association between weight change and suicide mortality, those having diabetes showed a J-shaped association between weight change and suicide mortality; especially, the group with severe weight gain showed 1.9-fold increased risk of suicide mortality. In subjects with type 2 diabetes, weight gain is associated with increased risk of medical illness such as cardiovascular disease [45], and contributes to patient frustration [46], which might increase the risk of suicide mortality.

The current study has several strengths. First, this study is a large-scale population-based cohort study including more than 2 million subjects. This enabled us to perform multiple subgroup analyses with statistical power. Second, the analyses were adjusted for the presence of chronic diseases such as diabetes, hypertension, dyslipidemia, and CKD, and especially the presence of depressive disorder. Nevertheless, this study also has some limitations. First, there is a possibility of selection bias, as the study only included individuals who participated in the general health screening program twice within a twoyear period. Those with higher health consciousness are more likely to engage in regular health screenings. Second, because this was an observational study, we could not exclude the possibility of unmeasured confounding factors contributing to the reverse J-shaped association between weight change and suicide mortality. Such factors can include conditions such as cardiovascular disease, heart failure, liver disease, chronic obstructive pulmonary disease, dementia, psychiatric disorders other than depression, as well as marital and employment status. Third, our analyses could not differentiate between intentional and unintentional weight loss. Because intentional weight loss to improve health can lower mortality [47, 48], this might affect the observed association between weight change and suicide mortality. In addition, this study did not include information on weight control behaviors, perceived weight status, suicidal ideation, and suicidal attempt, which were associated with suicide mortality [49, 50]. Fourth, we could not consider the onset time and duration of comorbidities, which might affect weight change. Fifth, because the diagnosis of depressive disorder was identified from claim data, there is a possibility of its underestimation. Sixth, this study did not include information on the use of psychiatric or antidiabetic medications that could have influenced weight changes. Lastly, the study results were derived from only Korean subjects. Therefore, the results cannot be generalized to other ethnicities.

## Conclusions

A reverse J-shaped association was found between weight change and suicide mortality; in particular, severe weight loss or weight gain showed 1.8-fold or 1.6-fold increased risk of suicide mortality, respectively. This association was consistently observed regardless of age, sex, depression, cancer, and BMI category. Our findings suggest that physicians should monitor subjects with severe weight change within a 2-year interval in relation to suicide behavior. Future studies incorporating information on weight change intentions, medications, weight changerelated medical conditions are needed to elucidate the underlying mechanisms through which weight change affects suicide mortality.

## Abbreviations

BP	Blood pressure
BMI	Body mass index
CVD	Cardiovascular disease
CKD	Chronic kidney disease
eGFR	Estimated glomerular filtration rate
HR	Hazard ratio
IR	Incidence rate
ICD-10	International Statistical Classification of Diseases, 10th revision
NHIS	National Health Insurance Service
Q	Quartile
WHO	World Health Organization

## **Supplementary Information**

The online version contains supplementary material available at https://doi.org/10.1186/s13098-024-01559-7.

Additional file 1.		
		)

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## Author contributions

K.K., J.H.J., Y.H.U., Y.B.A., S.H.K., K.H., and J.S.Y. were involved in the conception, design, and conduct of the study and the analysis and interpretation of the results. K.K., K.H., and J.S.Y. wrote the first draft of the manuscript, and all authors edited, reviewed, and approved the final version of the manuscript. K.H. and J.S.Y. are the guarantor of this work and, as such, had full access to all the data in the study.

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## Availability of data and materials

Data from the Korean National Health Insurance Service (NHIS) cannot be shared publicly due to the violation of patient privacy and the absence of informed consent for data sharing.

## Declarations

## Ethics approval and consent to participate

This study was approved by the Institutional Review Board of The Catholic University of Korea, St. Vincent's Hospital (VC23ZISE0218), and informed consent was waived. The study was conducted in accordance with the principles of the Declaration of Helsinki.

## **Consent for publication**

Not applicable.

### **Competing interests**

The authors declare no competing interests.

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## References

- Grundy SM. Obesity, metabolic syndrome, and cardiovascular disease. J Clin Endocrinol Metab. 2004;89(6):2595–600.
- Powell-Wiley TM, Poirier P, Burke LE, Despres JP, Gordon-Larsen P, Lavie CJ, Lear SA, Ndumele CE, Neeland IJ, Sanders P, et al. Obesity and cardiovascular disease: a scientific statement from the American heart association. Circulation. 2021;143(21):e984–1010.
- 3. Nuttall FQ. Body mass index: obesity, BMI, and health: a critical review. Nutr Today. 2015;50(3):117–28.
- Aune D, Sen A, Prasad M, Norat T, Janszky I, Tonstad S, Romundstad P, Vatten LJ. BMI and all cause mortality: systematic review and non-linear dose-response meta-analysis of 230 cohort studies with 3.74 million deaths among 30.3 million participants. BMJ. 2016;353:i2156.
- Bhaskaran K, Dos-Santos-Silva I, Leon DA, Douglas IJ, Smeeth L. Association of BMI with overall and cause-specific mortality: a populationbased cohort study of 3.6 million adults in the UK. Lancet Diabetes Endocrinol. 2018;6(12):944–53.
- Jee SH, Sull JW, Park J, Lee SY, Ohrr H, Guallar E, Samet JM. Bodymass index and mortality in Korean men and women. N Engl J Med. 2006;355(8):779–87.
- Corrada MM, Kawas CH, Mozaffar F, Paganini-Hill A. Association of body mass index and weight change with all-cause mortality in the elderly. Am J Epidemiol. 2006;163(10):938–49.
- Droyvold WB, Lund Nilsen TI, Lydersen S, Midthjell K, Nilsson PM, Nilsson JA, Holmen J, Nord-Trondelag Health S. Weight change and mortality: the nord-trondelag health study. J Intern Med. 2005;257(4):338–45.

- Adams KF, Leitzmann MF, Ballard-Barbash R, Albanes D, Harris TB, Hollenbeck A, Kipnis V. Body mass and weight change in adults in relation to mortality risk. Am J Epidemiol. 2014;179(2):135–44.
- 10. Lee IM, Paffenbarger RS Jr. Change in body weight and longevity. JAMA. 1992;268(15):2045–9.
- Kim YH, Kim SM, Han KD, Son JW, Lee SS, Oh SW, Lee WY, Yoo SJ. Change in weight and body mass index associated with all-cause mortality in Korea: a nationwide longitudinal study. J Clin Endocrinol Metab. 2017;102(11):4041–50.
- 12. World Health O. Suicide in the world: global health estimates. Geneva: World Health Organization; 2019.
- Garnett MF. Suicide mortality in the United States, 2001–2021. NCHS data brief, no 464. Hyattsville: National Center for Health Statistics; 2023.
- Jang H, Lee W, Kim YO, Kim H. Suicide rate and social environment characteristics in South Korea: the roles of socioeconomic, demographic, urbanicity, general health behaviors, and other environmental factors on suicide rate. BMC Public Health. 2022;22(1):410.
- Elovainio M, Shipley MJ, Ferrie JE, Gimeno D, Vahtera J, Marmot MG, Kivimaki M. Obesity, unexplained weight loss and suicide: the original Whitehall study. J Affect Disord. 2009;116(3):218–21.
- Kim B, Kim YC, Kim HS, Park SY, Choi JY. Significance of the regular publication of statistics on national health indicators in academic journals and the prospects of Korea national antimicrobial use analysis system (KONAS). Infect Chemother. 2023;55(2):306–7.
- Lee YH, Han K, Ko SH, Ko KS, Lee KU. Data analytic process of a nationwide population-based study using national health information database established by national health insurance service. Diabetes Metab J. 2016;40(1):79–82.
- Kim MK, Han K, Lee SH. Current trends of big data research using the Korean national health information database. Diabetes Metab J. 2022;46(4):552–63.
- Shin DW, Cho J, Park JH, Cho B. National general health screening program in Korea: history, current status, and future direction. Prec Future Med. 2022;6(1):9–31.
- Williamson DA, Bray GA, Ryan DH. Is 5% weight loss a satisfactory criterion to define clinically significant weight loss? Obesity. 2015;23(12):2319–20.
- 21. American College of Cardiology, American Heart Association Task Force on Practice Guidelines OEP. Executive summary: guidelines (2013) for the management of overweight and obesity in adults: a report of the American college of cardiology/American heart association task force on practice guidelines and the obesity society published by the obesity society and American college of cardiology/American heart association task force on practice guidelines. Based on a systematic review from the the obesity expert panel, 2013. Obesity. 2014;22(Suppl 2):S5-39.
- 22. Kim KK, Haam JH, Kim BT, Kim EM, Park JH, Rhee SY, Jeon E, Kang E, Nam GE, Koo HY, et al. Evaluation and treatment of obesity and its comorbidities: 2022 update of clinical practice guidelines for obesity by the Korean society for the study of obesity. J Obes Metab Syndr. 2023;32(1):1–24.
- 23. Pasman JA, Meijsen JJ, Haram M, Kowalec K, Harder A, Xiong Y, Nguyen TD, Jangmo A, Shorter JR, Bergstedt J, et al. Epidemiological overview of major depressive disorder in Scandinavia using nationwide registers. Lancet Reg Health Eur. 2023;29:100621.
- Jeong W, Kim H, Joo JH, Jang SI, Park EC. Association between depression and risk of Parkinson's disease in South Korean adults. J Affect Disord. 2021;292:75–80.
- Tyrrell J, Mulugeta A, Wood AR, Zhou A, Beaumont RN, Tuke MA, Jones SE, Ruth KS, Yaghootkar H, Sharp S, et al. Using genetics to understand the causal influence of higher BMI on depression. Int J Epidemiol. 2019;48(3):834–48.
- Yoo JE, Shin DW, Han K, Kim D, Jeong SM, Koo HY, Yu SJ, Park J, Choi KS. Association of the frequency and quantity of alcohol consumption with gastrointestinal cancer. JAMA Netw Open. 2021;4(8):e2120382.
- 27. Yoo JE, Han K, Shin DW, Kim D, Kim BS, Chun S, Jeon KH, Jung W, Park J, Park JH, et al. Association between changes in alcohol consumption and cancer risk. JAMA Netw Open. 2022;5(8):e2228544.
- Kwon S, Lee HJ, Han KD, Kim DH, Lee SP, Hwang IC, Yoon Y, Park JB, Lee H, Kwak S, et al. Association of physical activity with all-cause and cardiovascular mortality in 7666 adults with hypertrophic cardiomyopathy (HCM): more physical activity is better. Br J Sports Med. 2021;55(18):1034–40.
- Lee J, Jung J-H, Kang DW, Kim M-H, Lim D-J, Kwon H-S, Lee JM, Chang S-A, Han K, Lee S-H. Body weight variability and risk of suicide

mortality: a nationwide population-based study. Depress Anxiety. 2024;2024(1):7670729.

- Cerhan JR, Moore SC, Jacobs EJ, Kitahara CM, Rosenberg PS, Adami HO, Ebbert JO, English DR, Gapstur SM, Giles GG, et al. A pooled analysis of waist circumference and mortality in 650,000 adults. Mayo Clin Proc. 2014;89(3):335–45.
- Nanri A, Mizoue T, Takahashi Y, Noda M, Inoue M, Tsugane S. Weight change and all-cause, cancer and cardiovascular disease mortality in Japanese men and women: the Japan Public Health Center-Based Prospective Study. Int J Obes. 2010;34(2):348–56.
- Andres R, Muller DC, Sorkin JD. Long-term effects of change in body weight on all-cause mortality. A review. Ann Intern Med. 1993;119(7 Pt 2):737–43.
- Ahmedani BK, Simon GE, Stewart C, Beck A, Waitzfelder BE, Rossom R, Lynch F, Owen-Smith A, Hunkeler EM, Whiteside U, et al. Health care contacts in the year before suicide death. J Gen Intern Med. 2014;29(6):870–7.
- Han SH, Safeek R, Ockerman K, Trieu N, Mars P, Klenke A, Furnas H, Sorice-Virk S. Public interest in the off-label use of glucagon-like peptide 1 agonists (ozempic) for cosmetic weight loss: a google trends analysis. Aesthet Surg J. 2023;44(1):60–7.
- Alalwan AA, Friedman J, Park H, Segal R, Brumback BA, Hartzema AG. US national trends in bariatric surgery: a decade of study. Surgery. 2021;170(1):13–7.
- Jastreboff AM, Aronne LJ, Ahmad NN, Wharton S, Connery L, Alves B, Kiyosue A, Zhang S, Liu B, Bunck MC, et al. Tirzepatide once weekly for the treatment of obesity. N Engl J Med. 2022;387(3):205–16.
- Hecht LM, Yeh HH, Braciszewski JM, Miller-Matero LR, Thakrar A, Patel S, Simon GE, Lynch FL, Beck A, Owen-Smith AA, et al. Weighing the association between BMI change and suicide mortality. Psychiatr Serv. 2021;72(8):920–5.
- Otsuka T, Tomata Y, Sugawara Y, Tsuji I. Association between weight loss since the age of 20 years and the risk of suicide death: a populationbased cohort study. J Affect Disord. 2021;292:746–50.
- Vierboom YC, Preston SH, Stokes A. Patterns of weight change associated with disease diagnosis in a national sample. PLoS ONE. 2018;13(11):e0207795.
- 40. Ju YJ, Han KT, Lee TH, Kim W, Park JH, Park EC. Association between weight control failure and suicidal ideation in overweight and obese adults: a cross-sectional study. BMC Public Health. 2016;16:259.
- 41. Palmer BA, Pankratz VS, Bostwick JM. The lifetime risk of suicide in schizophrenia: a reexamination. Arch Gen Psychiatry. 2005;62(3):247–53.
- 42. Jamison KR. Suicide and bipolar disorder. J Clin Psychiatry. 2000;61(Suppl 9):47–51.
- 43. Ye W, Xing J, Yu Z, Hu X, Zhao Y. Mechanism and treatments of antipsychotic-induced weight gain. Int J Obes. 2023;47(6):423–33.
- Mangge H, Bengesser S, Dalkner N, Birner A, Fellendorf F, Platzer M, Queissner R, Pilz R, Maget A, Reininghaus B, et al. Weight gain during treatment of bipolar disorder (BD)-facts and therapeutic options. Front Nutr. 2019;6:76.
- Eeg-Olofsson K, Cederholm J, Nilsson PM, Zethelius B, Nunez L, Gudbjornsdottir S, Eliasson B. Risk of cardiovascular disease and mortality in overweight and obese patients with type 2 diabetes: an observational study in 13,087 patients. Diabetologia. 2009;52(1):65–73.
- Pi-Sunyer FX. The impact of weight gain on motivation, compliance, and metabolic control in patients with type 2 diabetes mellitus. Postgrad Med. 2009;121(5):94–107.
- French SA, Folsom AR, Jeffery RW, Williamson DF. Prospective study of intentionality of weight loss and mortality in older women: the Iowa Women's Health Study. Am J Epidemiol. 1999;149(6):504–14.
- Gregg EW, Gerzoff RB, Thompson TJ, Williamson DF. Intentional weight loss and death in overweight and obese U.S. adults 35 years of age and older. Ann Intern Med. 2003;138(5):383–9.
- Favril L, Yu R, Geddes JR, Fazel S. Individual-level risk factors for suicide mortality in the general population: an umbrella review. Lancet Public Health. 2023;8(11):e868–77.
- Eaton DK, Lowry R, Brener ND, Galuska DA, Crosby AE. Associations of body mass index and perceived weight with suicide ideation and suicide attempts among US high school students. Arch Pediatr Adolesc Med. 2005;159(6):513–9.

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