Body Mass Index and All-cause Mortality in a large Chinese Cohort

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1 ADSILACI	1	Abstract
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2	Background: Obesity, defined by the body mass index (BMI), is related to the
3	increased risk of death in Caucasians. However, the prospective association between
4	BMI and the risk of death in adult Taiwanese remains unclear.
5	Methods: We examined the association between BMI and all-cause mortality
6	prospectively among 58,738 men and 65,718 women, aged 20 years and older at
7	enrollment in 1998-1999, from four nationwide health screening centers in Taiwan.
8	Cox proportional hazards regression analyses were used to estimate the relative risks
9	of all-cause death for different BMI categories during a maximum follow-up of 10
10	years.
11	Results: 3,947 participants died during the follow-up period. The lowest risks of
12	death were observed at a BMI between 24.0 to 25.9 for both men and women (mean
13	BMI= 24.9 in both genders). After adjustment for age, smoking status, alcohol intake,
14	betel nut chewing, physical activity, income status, and education level, a U-shaped
15	association between BMI and all-cause death was observed. Similar U-shaped
16	relationship existed on different age groups, smoking status, having preexisting
17	chronic diseases, or if excluding deaths during the first 3 years of follow up.
18	Interpretation: The association between BMI and all-cause death was represented as
19	a U-shape relationship in adult Taiwanese. The lowest risk for all-cause death among
20	adult men and women was observed to relate to a BMI of 24.9.

Keywords: body mass index, mortality, obesity, underweight, Chinese

1 Introduction

2	The prevalence of obesity has dramatically increased in past decades in both
3	developed and developing countries. The World Health Organization (WHO) reported
4	that 1.6 billion adults are overweight and at least 400 million adults are obese; they
5	further predicted that by year 2015, approximately 2.3 billion adults will be
6	overweight and more than 700 million will be obese ¹ . In Taiwan, according to the
7	National Health and Nutrition Survey performed between 1993-1996 and 2005-2008,
8	the prevalence of overweight or obesity (defined as body mass index (BMI) \ge 24
9	kg/m ²) in men and women had also dramatically increased from 33.4 % and 31.7 % to
10	50.8 % and 36.9 % 2 . Overweight and obesity have been recognized as important and
11	independent risk factors for many chronic diseases, such as diabetes, hypertension,
12	stroke, cardiovascular diseases, and cancers ³⁻⁷ . Substantial epidemiological evidences
13	indicated that obesity is associated with increased cardiovascular and all-cause
14	mortality ^{8,9} . Therefore, obesity has become a major public health problem around the
15	world. Current definitions for obesity and overweight in adults were based on
16	Caucasian populations. The WHO has proposed another definition of obesity and
17	overweight for Asians, but most of these data were based on the cross-sectional
18	studies ¹⁰ . Previous study also found that, for a given BMI, Asians had higher body fat
19	than Caucasians ¹¹ . Furthermore, the association between BMI and all-cause mortality

1	had been found to be a J-shaped or U-shaped relationship. Most studies were done in
2	Caucasians, with only a few conducted amongst Asians. For example, Gu et al
3	reported that a U-shaped association existed between BMI and all-cause mortality in
4	Chinese ¹² . However, the participants in Gu's study were middle-aged adults with age
5	over 40 years, not general adults (age over 20 years). Therefore, we designed a large
6	prospective cohort study to assess the relationship between BMI and all-cause
7	mortality in a nationwide representative sample of Taiwanese adult (age over 20
8	years). We also intended to find the optimal BMI cut-off values for overweight or
9	obesity in Taiwanese adults.

1 Methods

2 Study subjects

3	The data were collected from four private nationwide MJ Health Screening
4	Centers in Taiwan from 1998 to 1999. Membership was required for attending the
5	programme. The registered health practitioners in these centers provide a
6	multidisciplinary team approach of health assessment programme for their members.
7	Most of them undergo health examination every 3-4 years and approximately 30 % of
8	them will receive the same health check-up every year. A total of 58,738 male and
9	65,718 female adults, aged 20 years and above, were recruited into the study. The
10	population structure in our study was similar to the national data of adult published by
11	the Taiwanese government. The population structure in our study was similar to the
12	national data of adult published by the Taiwanese government ¹³ . Deaths were
13	ascertained by computer linkage to the national death registry using ID number. All
14	deaths that occurred between study entry and December 2008 were included.
15	Anthropometric indices
16	Anthropometric characteristics were described in our previous report ^{14,15} . In
17	brief, trained staff measured height (measured to the nearest 0.1 cm) and weight
18	(measured to the nearest 0.1 kg) of each participant using an auto-anthropometer
19	(KN-5000A, Nakamura, Tokyo, Japan). Body mass index (BMI) was calculated as

1	weight (kg) divided by height squared (m^2). Study participants were grouped into 9
2	categories according to BMI at baseline (<18.5, 18.5-19.9, 20-21.9, 22-23.9, 24-25.9,
3	26-27.9, 28-29.9, 30-34.9, and \geq 35 kg/m ²). Approval for patient recruitment and data
4	analyses was obtained from the MJ Research Foundation Review Committee in
5	Taiwan. Informed consent was obtained from every subject. The reported
6	investigations were carried out in accordance with the principles of the Declaration of
7	Helsinki as revised in 2000.
8	Questionnaire
9	Cigarette smoking, alcohol intake, betel nut chewing, and physical activity
10	histories were recorded for each subject using questionnaires. Current, former, or
11	never users for smoking, alcohol intake, and betel nut chewing were defined as those
12	who reported the current use, any prior use, or never use of these substances,
13	respectively, at baseline survey. The cumulative exposure to smoking was assessed by
14	recording the duration (years) and quantity (number of cigarettes/day). Former users
15	were also asked for their age at quitting. Cumulative pack-years of smoking were
16	calculated as smoking-years multiplied by average daily cigarette use divided by 20.
17	Cumulative pack-years for smokers were categorized into two groups (Low: <10
18	pack-years; High: \geq 10 pack-years), so smoking status was categorized as none (0
19	pack-years), low (0~9.9 pack-years), and high (≥ 10 pack-years). Physical activity

1	was classified into three levels: none to mild (exercise less than one hour per week),
2	moderate (exercise one to four hours per week), and vigorous (exercise more than five
3	hours per week) physical activity. Income status was sub-divided into three levels:
4	low (< USD 12,500/year), middle (12,500-37,500/year), and high (>37,500/year).
5	Education was also sub-divided into three levels: low (elementary school and below),
6	middle (junior and senior high school), and high (college/university and above).
7	Participants who reported a history of stroke, cardiovascular disease, liver cirrhosis,
8	asthma, chronic renal disease, and cancer were defined as with preexisting chronic
9	diseases (n=15,180). Others were defined as health participants (n=109,276).
10	Statistical analysis
10	
11	The data are presented as the means and standard deviation for continuous
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11 12	The data are presented as the means and standard deviation for continuous variables. Analysis of variance (ANOVA) test was used to compare the continuous
11 12 13	The data are presented as the means and standard deviation for continuous variables. Analysis of variance (ANOVA) test was used to compare the continuous variables (such as age, height, weight) across BMI groups in Table 1. Proportions and
11 12 13 14	The data are presented as the means and standard deviation for continuous variables. Analysis of variance (ANOVA) test was used to compare the continuous variables (such as age, height, weight) across BMI groups in Table 1. Proportions and categorical variables (such as smoking, alcohol drinking, and betel nut chewing status)
 11 12 13 14 15 	The data are presented as the means and standard deviation for continuous variables. Analysis of variance (ANOVA) test was used to compare the continuous variables (such as age, height, weight) across BMI groups in Table 1. Proportions and categorical variables (such as smoking, alcohol drinking, and betel nut chewing status) were tested by the χ^2 test and by the two-tailed Fisher's exact method when
 11 12 13 14 15 16 	The data are presented as the means and standard deviation for continuous variables. Analysis of variance (ANOVA) test was used to compare the continuous variables (such as age, height, weight) across BMI groups in Table 1. Proportions and categorical variables (such as smoking, alcohol drinking, and betel nut chewing status) were tested by the χ^2 test and by the two-tailed Fisher's exact method when appropriate in Table 1. Cox proportional hazards regression analyses adjusted for

- 1 statistical analyses were performed using the PC version of SPSS statistical software
- 2 (13th version, SPSS Inc., Chicago, IL, USA).

Results

2	During a maximum follow-up of 10 years (1,109,273 person-years), 2,398 men
3	and 1,549 women died. In Table 1, we showed the baseline characteristics of study
4	participants across 9 BMI categories. Participants with a higher BMI, as compared
5	with those who had a lower BMI, were characterized by older age, more likely to
6	consume alcohol and betel nut chewing currently, less physical activity, with higher
7	income status in men and a lower income status and lower educations level in
8	women,.
9	In Table 2 , we showed the RRs for the risk of all-cause death among men and
10	women. It poses a nonlinear association between BMI and the risk of all-cause death.
11	The lowest risk in relation to all-cause death among both male and female participants
12	was found at a BMI between 24.0 to 25.9 kg/m ² . There was a statistically significant
13	U-shaped association between BMI and all-cause death. Using participants with a
14	BMI between 24.0 to 25.9 as the reference group, and with the Cox proportional
15	hazards regression analyses with adjustment for age, smoking status, alcohol intake,
16	betel nut chewing, physical activity, income status, and education level, the RRs for
17	all-cause death across BMI categories among men and women were showed in Table
18	2 and Figure 1.

19 There were significant interactions (P < 0.05) between BMI categories and

1	preexisting chronic disease status/age group/smoking status for predicting the risk of
2	all-cause mortality. We therefore stratified these groups and presented the results in
3	Figure 2. Figure 2 showed the adjusted RRs in relation to all-cause death according
4	to smoking status (never smoker, low-dose smokers, and high-dose smokers; Figure
5	2a), different age groups (participants aged 20 to 64 years old and aged 65 years or
6	more; Figure 2b), and with or without preexisting chronic diseases (Figure 2c). The
7	U-shaped association between BMI and all-cause death persisted within these three
8	smoking groups and within the two age groups. High-dose and low-dose smokers
9	seem to have higher RRs for all-cause mortality in relation to BMI than the never
10	smokers with increased risks in the lower BMI categories. In comparison to the
11	high-dose smokers, for those participants in the higher BMI categories, the RRs for
12	all-cause mortality were higher among never and low-dose smokers. In comparison to
13	participants less than 65 years, the RRs for all-cause mortality in lower and higher
14	BMI categories were higher among those with age over 65 years. The lowest all-cause
15	mortality was also found in BMI between 24.0 to 25.9 in both age groups. Participants
16	with or without preexisting chronic diseases also exhibited the U-shaped relationship
17	between BMI and all-cause death. Participants with preexisting chronic diseases in the
18	lower BMI categories had higher risks of death than healthy participants. In contrast,
19	participants with preexisting chronic diseases in the higher BMI categories had lower

1 risks of death than healthy participants.

2	After excluding study participants who died during the first 3 years of follow-up
3	and using BMI between 24.0 to 25.9 for comparison, the adjusted RRs for all-cause
4	death across BMI categories were also showed in Table 2. The U-shaped association
5	between BMI and all-cause death still existed among both men and women.
6	

1 Interpretation

2	In this population-based prospective study, we demonstrated that the higher risks of
3	death were observed in the lower and upper BMI categories compared to the middle
4	categories, showing a U-shaped relationship between BMI and all-cause mortality in
5	adult Taiwanese. The same U-shaped association existed on those who were either
6	aged over 65 or less than 65 years old, or who were never smokers, low-dose smokers,
7	or high-dose smokers, or who were with or without preexisting chronic diseases, or
8	who did not die during the first 3 years of follow-up. In addition, we also found that
9	the lowest death in relation to BMI was observed at a BMI of 24.9 for both men and
10	women.
11	The WHO has proposed a lower BMI cut-off values for defining overweight and
12	obesity in Asians. However, most of this evidence was based on cross-sectional
13	studies. Some prospective studies done in Chinese have reported the U-shaped
14	relationship between BMI and all-cause mortality and found subjects with BMI
15	between 24.0 to 28.0 have the lowest risks of all-cause death ^{12,16,17} . For example, Gu
16	et al reported that subjects with BMI between 24.0 to 24.9 in both men and women
17	had the lowest risk of all-cause death in 169,871 Chinese aged 40 years and older.
18	Studies done in Caucasians have also identified a J- or U-shaped association between
	Studies done in Caucasians have also identified a 3- of O-shaped association between

1	23.0 to 28.0 had lowest risk in relation to all-cause mortality and the minimum
2	mortality close to a BMI of 25.0 ^{8,18-20} . Our findings are consistent with these studies
3	and do not support the use of a lower BMI cut-off value for obesity in Taiwanese
4	general adults. We also conducted the sensitivity analyses to demonstrate the optimal
5	cut-off values of BMI for all-cause mortality by dividing BMI into 15 groups
6	according to BMI- <18.5, 18.5-19.9, 20-20.9, 21-21.9, 22-22.9, 23-23.9, 24-24.9,
7	25-25.9, 26-26.9, 27-27.9, 28-28.9, 29-29.9, 30-34.9, \geq 35 kg/m ² . The lowest risk of
8	death was observed at a BMI of 24.0 to 24.9 for men and 25.0 to 25.9 for women
9	(mean BMI= 24.5 in men and 25.5 in women) which were also consistent with our
10	results (BMI: 24.0-25.9). These findings demonstrated the robustness of our results.
11	Previous studies had found that age was an effect modifier for the relationship
12	between BMI and the risk of death ^{18,20} . For example, Weiss et al reported that BMI is
13	inversely related to mortality in the elderly ²¹ . Our study, however, found that the
14	similar U-shaped association between BMI and all-cause death was presented among
15	participants aged 20 to 64 years and among those 65 years and above. This finding did
16	not support the theory of "obesity paradox" in the elderly.
17	Smoking is associated with lower body weight and an increased risk of death ²²⁻²⁴ .
18	Compared to previous studies ^{12,23} , our findings are consistent with the findings that
19	among current smokers, subjects in lower BMI categories had higher risk of death.

1	However, among never smokers we did not find subjects in the higher BMI categories
2	had higher risk of death. Similar U-shaped relationships between BMI and all-cause
3	mortality were found among never smokers, low-dose smokers, and high-dose
4	smokers.
5	Preexisting chronic diseases were linked to an increase in all-cause death and
6	decrease in body weight. Two criteria were used to avoid possible bias caused by
7	preexisting chronic diseases: 1) limited the analysis to healthy participants and 2)
8	excluded those who died during the first 3 years of follow-up. In this study, similar
9	U-shaped association between BMI categories and all-cause death remained existed
10	after making these adjustments. In the lower BMI categories, the risk of death was
11	higher for those participants with preexisting chronic diseases, compared to healthy
12	participants. This confirmed that preexisting chronic diseases were associated with the
13	increase of all-cause death. In contrary, in the higher BMI categories, the risk of death
14	among participants with preexisting chronic diseases was lower compared to the
15	healthy participants. This suggested that obesity may be a protective factor among
16	subjects with preexisting chronic diseases. However this finding was consistent with
17	the reports from other populations i.e. see Adams et al. in the United States ²⁵ .
18	Limitations

Although we have shown that a U-shaped relationship existed between BMI and

1	all-cause mortality in adult Taiwanese, there are some limitations in this study. Firstly,
2	we didn't measure the weight at the end of follow-up, so we could not get information
3	about weight changes over time. Hence, we could not find the relationship between
4	weight change and mortality. Secondly, our study population mainly came from
5	generally healthy volunteers who attended health screening centers rather than
6	nationally representative subjects. However, the population structure in our study was
7	similar to the national data of adult published by the Taiwanese government. External
8	validation will be necessary in future studies. Finally, we have adjusted several
9	variables, including smoking status, alcohol intake, betel nut chewing, physical
10	activity, level of education, and level of income, which allowed us to minimize the
11	potential confounders. However, we still could not exclude the possibility of the
12	residual confounding.
13	Conclusion
14	We have demonstrated that both obesity and underweight are related to increase
15	all-cause mortality in adult Taiwanese population. This relation was found in persons
16	who were above or less than 65 years old, were heavy, mild, or never smokers, were
17	with or without preexisting chronic diseases, and who did not die during the first 3
18	years of follow-up. The consistency of our findings in both men and women and
19	across all ages (younger to middle-aged adults and in the elderly) also argues for the

1 use of a single recommended range of body weight throughout life.

1 **Competing interests:** None declared

2	Contributions: Dr. WY Lin and KC Huang had full access to all of the data in the
3	study and take responsibility for the integrity of the data and the accuracy of the data
4	analysis. Study concept and design: Dr. WY Lin, CCLin, and KC Huang. Acquisition
5	of data: Dr. PK Song, SL Tsai, and TC Li. Analysis and interpretation of data: Dr. WY
6	Lin, JB Albu, FX Pi-Sunyer, and TC Li. Drafting of the manuscript: Dr. WY Lin and
7	KC Huang. Critical revision of the manuscript for important intellectual content: Dr.
8	JB Albu, FX Pi-Sunyer, CC Lin, TC Li, and KC Huang. Statistical analysis: Dr. WY
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REFERENCE

1. World Health Organization. Obesity and overweight. 2006;Fact sheet

311:http://www.who.int/mediacentre/factsheets/fs311/en/index.html. Accessed at 10, Dec., 2010.

2. Department of Health, Executive Yuan, TAIWAN. Prevalence of overweight, obesity, and metabolic syndrome in Taiwan. 2009: http://www.doh.gov.tw/EN2006/index_EN.aspx. Accessed at 10, Dec., 2010.

3. Calle EE. Obesity and cancer. *BMJ* 2007;335:1107-8.

4. World Health Organization. Obesity: preventing and managing the global epidemic. Report of a WHO consultation. *WHO Tech Rep Ser* 2000;894:1-253.

 Narayan KM, Boyle JP, Thompson TJ, et al. Effect of BMI on lifetime risk for diabetes in the U.S. Diabetes Care 2007;30:1562-6.

6. Lin WY, Yang WS, Lee LT, et al. Insulin resistance, obesity, and metabolic syndrome among non-diabetic pre- and post-menopausal women in North Taiwan. *Int J Obes (Lond)* 2006;30:912-7.

7. Lin WY, Yao CA, Wang HC, et al. Impaired lung function is associated with obesity and metabolic syndrome in adults. *Obesity (Silver Spring)* 2006;14:1654-61.

Pischon T, Boeing H, Hoffmann K, et al. General and abdominal adiposity and risk of death in Europe.
 N Engl J Med 2008;359:2105-20.

9. Kannel WB, D'Agostino RB, Cobb JL. Effect of weight on cardiovascular disease. *Am J Clin Nutr* 1996;63:419S-22S.

World Health Organization. The Asia-Pacific perspective:Redefining obesity and its treatment.
 WHO:Geneva. 2000. Accessed at 10, Dec., 2010.

11. Chang CJ, Wu CH, Chang CS, et al. Low body mass index but high percent body fat in Taiwanese

subjects: implications of obesity cutoffs. Int J Obes Relat Metab Disord 2003;27:253-9.

12. Gu D, He J, Duan X, et al. Body weight and mortality among men and women in China. *JAMA* 2006;295:776-83.

Department of Health, Executive Yuan, Taiwan. Taiwan Public Health Report 1998-2000. DOH: Taipei,.
 2001:http://www.doh.gov.tw.

14. Lin WY, Chiu TY, Lee LT, et al. Betel nut chewing is associated with increased risk of cardiovascular disease and all-cause mortality in Taiwanese men. *Am J Clin Nutr* 2008;87:1204-11.

15. Lin WY, Lee LT, Chen CY, et al. Optimal cut-off values for obesity: using simple anthropometric

indices to predict cardiovascular risk factors in Taiwan. Int J Obes Relat Metab Disord 2002;26:1232-8.

16. Zhou BF. Effect of body mass index on all-cause mortality and incidence of cardiovascular diseases--report for meta-analysis of prospective studies open optimal cut-off points of body mass index in Chinese adults. *Biomed Environ Sci* 2002;15:245-52.

17. Yuan JM, Ross RK, Gao YT, et al. Body weight and mortality: a prospective evaluation in a cohort of middle-aged men in Shanghai, China. *Int J Epidemiol* 1998;27:824-32.

18. Flegal KM, Graubard BI, Williamson DF, et al. Excess deaths associated with underweight, overweight, and obesity. *JAMA* 2005;293:1861-7.

19. Allison DB, Fontaine KR, Manson JE, et al. Annual deaths attributable to obesity in the United States. *JAMA* 1999;282:1530-8.

20. Calle EE, Thun MJ, Petrelli JM, et al. Body-mass index and mortality in a prospective cohort of U.S. adults. *N Engl J Med* 1999;341:1097-105.

21. Weiss A, Beloosesky Y, Boaz M, et al. Body mass index is inversely related to mortality in elderly

subjects. J Gen Intern Med 2008;23:19-24.

22. Gu D, Kelly TN, Wu X, et al. Mortality attributable to smoking in China. N Engl J Med 2009;360:150-9.

23. Miyazaki M, Babazono A, Ishii T, et al. Effects of low body mass index and smoking on all-cause mortality among middle-aged and elderly Japanese. *J Epidemiol* 2002;12:40-4.

24. Wilson K, Clark H, Hotz S, et al. Impact of smoking status on weight loss and cardiovascular risk factors. *J Epidemiol Community Health* 2001;55:213-4.

25. Adams KF, Schatzkin A, Harris TB, et al. Overweight, obesity, and mortality in a large prospective cohort of persons 50 to 71 years old. *N Engl J Med* 2006;355:763-78.

Table 1. Baseline Characteristics of Participants according to BMI among Men and Women¹

	<18.5	18.5-19.9	20-21.9	22-23.9	24-25.9	26-27.9	28-29.9	30-34.9	≥35
Men (N)	2448	4300	10473	14983	13368	7738	3374	1819	235
Age (years) ^{2,4,5}	39.3(15.6)	39.7(14.8)	41.1(14.5)	43.3(13.8)	44.6(13.4)	45.1(13.3)	44.4(13.4)	41.7(12.9)	37.5(11.9)
Height (cm) ^{2,4,5}	169.1(6.6)	169.0(6.5)	168.8(6.2)	168.5(6.3)	168.2(6.2)	168.1(6.1)	168.4(6.3)	169.0(6.4)	170.2(7.1)
Weight (kg) ^{2,4,5}	50.3(4.6)	55.3(4.4)	60.1(4.7)	65.4(5.1)	70.4(5.4)	76.1(5.8)	81.9(6.4)	90.3(8.0)	109(11.4)
BMI (kg/m ²) ^{2,4,5}	17.5(0.8)	19.3(0.4)	21.1(0.6)	23.0(0.6)	24.9(0.6)	26.9(0.6)	28.8(0.6)	31.6(1.3)	37.7(3.9)
Smoking (%) ^{3,5}									
Current	46.2	43.6	40.7	38.7	39.4	40.5	42.5	44.7	56.5
Former	9.7	9.7	10.6	12.1	13.3	13.5	14.3	13.0	8.1
Never	44.1	46.7	48.7	49.2	47.3	46.0	43.2	42.3	35.4
Alcohol intake (%) ^{3,5}									
Current	24.3	25.0	27.7	30.2	33.0	33.9	34.4	35.1	22.8
Former	5.8	5.0	5.2	5.0	5.4	6.3	7.0	6.7	7.8
Never	69.9	70.0	67.1	64.8	61.6	59.8	58.6	58.2	69.4
Betel nut chewing (%) ^{3,5}									
Current	9.6	10.1	9.5	9.7	10.6	12.2	13.8	15.6	18.8
Former	10.9	9.5	9.3	8.5	9.8	11.0	12.6	14.8	18.4
Never	79.5	80.4	81.2	81.8	79.6	76.8	73.7	69.6	62.8

Body mass index (BMI, kg/m²)

Physical activity (%) ^{3,5}									
None/mild	52.1	48.6	45.5	42.6	42.8	44.7	46.4	51.0	55.6
Moderate	34.9	37.5	38.4	39.4	38.5	37.2	36.5	34.5	34.7
Vigorous	13.1	13.8	16.1	18.0	18.7	18.1	17.1	14.5	9.7
Income (%) ^{3,5}									
Low	39.7	37.2	32.2	27.8	27.0	28.3	29.3	29.0	34.6
Moderate	55.1	57.1	59.5	61.5	60.6	59.2	58.5	60.3	55.1
High	5.2	5.7	8.3	10.7	12.4	12.5	12.2	10.7	10.3
Education (%) ^{3,5}									
Low	13.5	13.7	13.5	15.5	16.9	18.3	18.9	15.8	12.6
Moderate	38.2	36.1	34.6	33.2	34.8	36.1	37.5	39.0	41.7
High	48.3	50.2	51.8	51.3	48.4	45.6	43.6	45.2	45.7
Women (N)	6850	10171	16054	13253	9224	5322	2633	1908	303
Age (years) ^{2,4,5}	32.7(10.6)	35.5(11.1)	40.1(12.2)	45.5(13.1)	49.5(12.8)	51.6(12.5)	51.8(12.4)	51.2(13.0)	47.6(13.5)
Height (cm) ^{2,4,5}	158.6(5.6)	158.0(5.5)	157.0(5.6)	155.7(5.6)	155.0(5.5)	154.4(5.5)	154.3(5.6)	154.3(5.6)	154.4(6.3)
Weight (kg) ^{2,4,5}	44.0(3.7)	48.3(3.5)	51.8(3.8)	55.7(4.1)	59.9(4.4)	64.2(4.7)	68.9(5.1)	75.6(6.3)	89.7(9.5)
BMI (kg/m ²) ^{2,4,5}	17.5(0.8)	19.3(0.4)	21.0(0.6)	23.0(0.6)	24.9(0.6)	26.9(0.6)	28.9(0.6)	31.7(1.3)	37.6(2.4)
Smoking (%) ^{3,5}									
Current	9.0	7.3	5.6	4.7	3.9	4.2	4.3	5.0	6.7
Former	2.2	2.0	1.6	1.3	1.1	1.0	1.5	1.5	2.1

Never	88.8	90.7	92.8	94.0	95.0	94.8	94.2	93.5	91.2
Alcohol intake (%) ^{3,5}									
Current	5.6	4.9	5.9	5.7	5.7	5.6	4.6	5.3	6.2
Former	1.5	1.3	1.4	1.6	1.5	1.4	1.3	2.3	6.2
Never	92.9	93.8	92.7	92.8	92.8	92.9	94.1	92.4	87.6
Betel nut chewing (%) ^{3,5}									
Current	0.2	0.3	0.3	0.3	0.5	0.9	1.3	1.6	2.9
Former	0.2	0.1	0.2	0.4	0.3	0.5	0.5	0.9	0.4
Never	99.6	99.6	99.5	99.3	99.2	98.6	98.2	98.5	96.7
Physical activity (%) ^{3,5}									
None/mild	67.2	60.2	54.7	50.2	49.0	48.9	52.7	54.6	59.2
Moderate	28.1	32.9	35.1	34.3	32.8	31.6	30.0	29.0	29.6
Vigorous	4.7	7.0	10.2	15.5	18.2	19.5	17.2	16.5	11.2
Income (%) ^{3,5}									
Low	57.8	54.3	57.0	65.6	73.4	78.5	79.4	82.8	80.1
Moderate	39.8	42.7	39.6	31.6	24.4	19.5	17.9	15.7	18.1
High	2.4	3.0	3.4	2.8	2.2	2.1	2.7	1.5	1.8
Education (%) ^{3,5}									
Low	7.0	9.9	18.6	34.7	48.5	58.5	61.6	62.0	56.7
Moderate	38.6	39.7	40.5	37.0	32.8	28.5	26.6	26.1	28.9

High	54.4	50.5	40.9	28.3	18.7	13.0	11.8	11.9	14.4

¹ BMI, body mass index;

² ANOVA test was used for comparing mean values of continuous variables between groups

³ Pearson chi-square test was used for categorical data

⁴ mean (SD) (all such values).

⁵ p values < 0.001

	<18.5	18.5-19.9	20-21.9	22-23.9	24-25.9	26-27.9	28-29.9	30-34.9	≥35
Men									
Deaths (no.)	145	196	425	586	506	310	148	71	11
Person-years	21507	38092	92829	133037	118723	68759	29954	16083	2080
Relative risk (9	5%CI)								
Crude ¹	1.80(1.50-2.17)	1.47(1.24-1.73)	1.26(1.11-1.44)	1.11(0.98-1.25)	1.00	1.03(0.90-1.19)	1.18(0.98-1.42)	1.40(1.09-1.80)	2.48(1.36-4.50)
Adjusted ²	1.65(1.34-2.04)	1.32(1.09-1.60)	1.15(0.99-1.33)	1.14(0.99-1.30)	1.00	1.01(0.86-1.18)	1.12(0.91-1.38)	1.27(0.95-1.70)	2.37(1.22-4.60)
Follow-up perio	od (1990-2007) ³								
Adjusted ²	1.65(1.29-2.10)	1.36(1.10-1.69)	1.10(0.92-1.30)	1.09(0.94-1.27)	1.00	1.00(0.84-1.20)	1.19(0.95-1.50)	1.09(0.76-1.55)	2.08(0.92-4.67)
Women									
Deaths (no.)	90	128	265	332	306	202	117	95	14
Person-years	61448	91146	143926	118733	82357	47488	23422	16989	2698
Relative risk (9	5%CI)								
Crude ¹	1.50(1.19-1.90)	1.22(0.99-1.50)	1.05(0.89-1.24)	0.99(0.85-1.16)	1.00	1.00(0.84-1.20)	1.16(0.94-1.44)	1.34(1.07-1.69)	1.63(0.96-2.79)
Adjusted ²	1.52(1.17-1.99)	1.34(1.06-1.70)	1.18(0.97-1.42)	1.05(0.88-1.26)	1.00	1.03(0.84-1.26)	1.08(0.83-1.39)	1.39(1.07-1.80)	1.65(0.92-2.95)
Follow-up perio	od (1990-2007) ³								
Adjusted ²	1.60(1.19-2.15)	1.22(0.93-1.61)	1.22(0.99-1.51)	1.03(0.84-1.26)	1.00	1.07(0.85-1.34)	1.03(0.77-1.38)	1.44(1.08-1.93)	1.76(0.93-3.34)

Body Mass Index (BMI, kg/m²)

¹Crude: adjusted for age.

² Adjusted 1: adjusted for age, smoking status (current, former, never), alcohol intake, betel nut chewing,

physical activity, income, and education level.

³Excluded those who died during first 3 years follow-up (only deaths that occurred after more than 3 years of follow-up were included).

Legend to Figure 1

Adjusted relative risks of all-cause mortality according to BMI categories among men and women. Relative risks adjusted for age, smoking status, alcohol intake, betel nut chewing, physical activity, income, and education level (a) in men; (b) in women. Error bars indicate 95 % confidence interval.

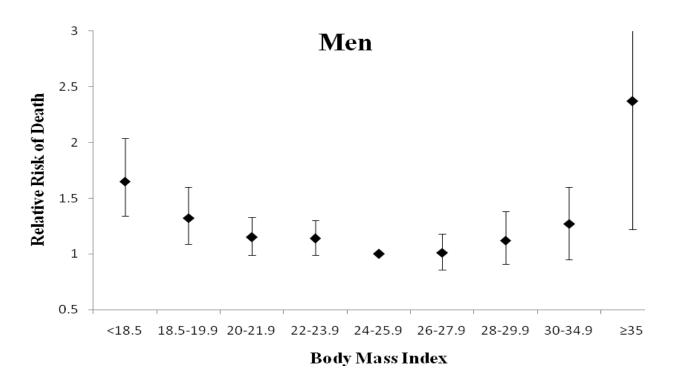
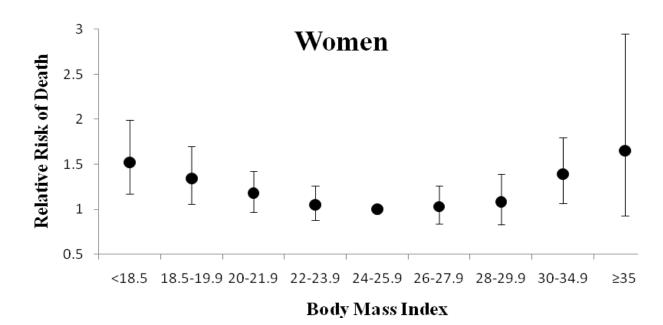


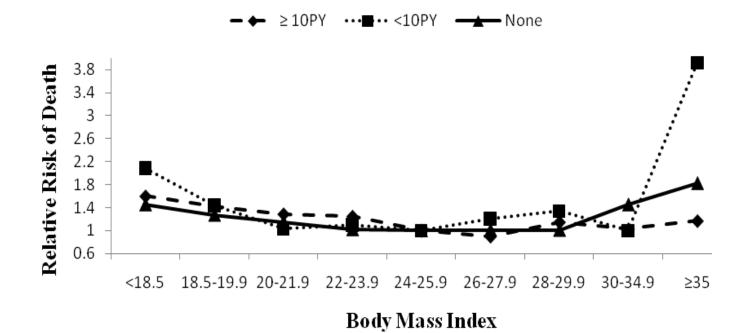
Figure 1 (b).



Legend to Figure 2

Adjusted relative risks of all-cause mortality according to BMI categories by smoking status, age group, and presence of preexisting chronic diseases. (a) Smoking status was divided into three categories by the consumption and smoking duration: none, <10 PY, and \geq 10 PY (PY: pack per day x smoking duration (year)). The relative risks were adjusted for age, gender, cumulative smoking amount, alcohol intake, betel nut chewing, income, and education level; (b) age was divided into two categories: <65 years old and \geq 65 years old; (c) participants with and without preexisting chronic diseases (previous history of stroke, cardiovascular disease, liver cirrhosis, chronic renal disease, asthma, or cancers). The relative risks were adjusted for age, gender, smoking status, alcohol intake, betel nut chewing, physical activity, income, and education level.

Figure (2a).



32

