# Peer

## Depressive symptoms and other risk factors predicting suicide in middle-aged men: a prospective cohort study among Korean Vietnam War veterans

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### ABSTRACT

**Background.** Few studies have prospectively examined whether depressive symptoms and other risk factors are associated with a higher risk of suicide death in individuals other than high-risk populations such as psychiatric patients and individuals with self-harm histories. The purpose of the study is to prospectively examine whether depressive symptoms assessed by the Beck Depression Inventory (BDI) are associated with greater risk of suicide death and whether depressive symptoms and other risk factors are independent predictors of suicide in general-risk populations. Another aim is to evaluate the sensitivity of the BDI for predicting suicide death.

**Methods.** 10,238 Korean Vietnam War veterans (mean age: 56.3 years) who participated in two surveys in 2001 were followed up for suicide mortality over 7.5 years.

**Results.** 41 men died by suicide. Severely depressed participants had a higher adjusted hazard ratio (aHR = 3.4; 95% CI [1.5–7.7]) of suicide than non-to-moderately depressed ones. Higher suicide risk was associated with more severe depressive symptoms (*p* for trend = 0.009). After adjustment for depressive symptoms and other factors, very poor health, low education, and past drinking were associated with higher suicide risk, while good health, body mass index, and marital status were not associated with suicide. The sensitivity at the cut-off score of 31 for detecting suicide was higher during the earlier 3.5 years of the follow-up (75%; 95% CI [50–90]) than during the latter 4 years (60%; 95% CI [41–76]).

**Conclusions.** Depressive symptoms are a strong independent predictor and very poor health, low education, and drinking status may be independent predictors of future suicide. The BDI may have acceptable diagnostic properties as a risk assessment tool for identifying people with depression and suicidal potential among middle-aged men.

**Subjects** Epidemiology, Psychiatry and Psychology, Public Health **Keywords** Asia, Beck Depression Inventory, Cohort studies, Men, Depressive symptoms, Suicide

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#### INTRODUCTION

Over 800,000 people die by suicide worldwide every year (World Health Organization, 2014). Suicide prevention is a high priority in many countries. Understanding the risk factors leading to suicide is crucial for effectively preventing suicide and managing individuals at risk of suicide. Depression has been strongly associated with suicide in retrospective psychological autopsy studies (Hawton & van Heeringen, 2009; Yoshimasu, Kiyohara & Miyashita, 2008), although an over-underestimation of the association, and biases such as interviewer bias, surrogate interview bias, and recall bias may not be ruled out (Hielmeland et al., 2012). Prospective studies have been conducted mainly among patients with psychiatric disorders, who may not represent the people with psychiatric disorders in the community (Agerbo, 2007). Meanwhile, physical health indicators such as health status (Jee et al., 2011) and body mass index (BMI) (Jee et al., 2011; Mukamal et al., 2010), marital status (Masocco et al., 2010), drinking status (Akechi et al., 2006), and socioeconomic status (Milner, Page & Lamontagne, 2014) have also been linked with suicide. Despite many studies supporting these associations, it has seldom been evaluated and is unclear whether each of those factors were independently associated with suicide after adjusting for the others and, notably, for depression, one of the most important known precursors of suicide (Milner, Page & Lamontagne, 2014; Mukamal et al., 2010).

Meanwhile, suicide risk assessment using depression tools could be relatively easily embedded in the various health care settings to help primary physicians and public health professionals identify people with depression and suicidal potential. However, to the best of our knowledge, the diagnostic characteristics of depression instruments that can be used for suicide risk assessment have never been evaluated in general-risk populations for predicting suicide death.

The primary aim of this study was to prospectively examine whether more severe depressive symptoms assessed by the Korean translation of the Beck Depression Inventory (BDI) would be associated with a greater risk of suicide in middle-aged Korean men, specifically Korean veterans of the Vietnam War. The secondary aim was to evaluate the diagnostic properties of the BDI for predicting suicide death, such as sensitivity, specificity, and area under the receiver operating characteristics (ROC) curve (AUC). The tertiary aim was to examine whether risk factors such as self-rated health status, drinking, BMI, marital status, and socio-economic status were independently associated with suicide after adjusting for each other and for depressive symptoms. Korean Vietnam War veterans have had the same suicide mortality (crude mortality, 35.4 deaths per 100,000 person-years during 1992–2005; standardized mortality ratio = 0.95, 95% CI [0.89–1.02]) expected from the general Korean population in previous research (*Yi & Ohrr, 2011*).

#### **METHODS**

#### **Study participants**

We identified 187,897 Vietnam War veterans during 1999–2000 with the cooperation of the Ministry of Defense and then the Ministry of Government Administration and Home Affairs (*Yi et al., 2014*). Among them, 20,000 veterans were selected by a simple random

sample. After excluding 1,737 individuals who were deceased, had emigrated to another country, or had an unknown residency status as of July 2000, 18,263 were recruited for a health exam or survey. Among them, randomly selected 5,000 veterans (health exam group) were invited to a health exam and a survey by mail on May 4, 2001, and the other 13,263 (survey-only group) received a survey sent out on July 20, 2001. In the end, 10,238 veterans who participated in the health exam or survey (health exam group, n = 2,005; survey-only group, n = 8,233) were followed up on suicide mortality until December 31, 2008 (Fig. S1). The veterans were provided with a written summary about the research and an invitation to the health exam by post. Return of the postal survey and voluntary participation of health exam were considered implied consent. This study was approved by the Institutional Review Board of Kwandong University (Approval No: KD-13-0105).

#### Follow-up and ascertainment of suicide death

The deaths from suicide were ascertained from the 2001–2008 death records of the National Statistical Office. When veterans had emigrated to another country or their residential status was unknown between enrollment and December 2008, the changed date of the residential status was considered the date of a loss to follow-up. A complete follow-up was made for 10,181 veterans (99.4%). Suicide was identified by the International Classification of Diseases, 10th Revision codes X60-X84.

#### Measures of depressive symptoms

The self-administered BDI, with 21 items each measured on a 4-point Likert scale (0–3) and a total score range of 0–63, was used to assess depressive symptoms (*Beck et al.*, 1961). The previous Korean-language version of the BDI, translated by two psychiatrists and a clinical psychologist (*Rhee et al.*, 1995), was modified for the present study by two medical doctors, and several graduate students of public health. The internal consistency assessed by Cronbach's alpha was 0.937 in a complete data analysis.

The participants were categorized into two groups (no-to-moderate [0–30] and severe [31–63]) and five groups (depressive symptoms [BDI score]: none [0–13], mild [14–21], moderate [22–30], severe [31–39], and extreme [40–63]) based on quartiles and the highest decile of BDI score. For a sensitivity analysis, they were also categorized based on cut-off score in previous research into two groups (no-to-moderate [0–29] and severe [30–63]) and five groups (none [0–9], mild [10–16], moderate [17–29], severe [30–40], and extreme [41–63]) (*Al-Turkait & Ohaeri, 2010; Beck & Steer, 1987; Smarr & Keefer, 2011*).

#### Measures of other risk factors

Self-rated health, smoking, drinking ("Do you drink alcohol?" [Yes, No. I quit drinking, or Never]), marital status, educational status, and household monthly income were collected through self-reported survey. Self-rated health was assessed by a 5-point Likert scale. Weight and height were measured among participants in health exams, while they were self-reported among the survey-only group. The BMI was obtained from weight divided by the height squared (kg/m<sup>2</sup>).

#### Statistical analysis

Among the 10,238 participants, 3,156 (30.8%) had one or more missing baseline data including smoking (n = 55), drinking (n = 84), BMI (n = 102), self-rated health (n = 186), marital status (n = 143), educational status (n = 195), income status (n = 426), or at least one item on the BDI (n = 2761). Multiple imputation was performed to estimate the missing baseline data with SAS PROC MI. Markov chain Monte Carlo sampling methods and an expectation–maximization algorithm with a separate chain for each imputation (chain = multiple) were used. 100 imputations were made using all variables analyzed in this study and other variables including military variables (such as Agent Orange exposure index, unit served and military rank during Vietnam service), and self-rated health compared to the same age group. Imputation methods including number of imputations were selected to minimize imputation variance and to ensure the confidence intervals and p values to be reliable (*White, Royston & Wood, 2011*).

The chi-squared test and analysis of variance were performed to compare individual characteristics by two categories of depressive symptoms. A Cox proportional hazards model including various risk factors was implemented to calculate hazard ratios (HRs). The variables included in the Cox model were age at enrollment (years), smoking (current, past, and never smoker), drinking (current, past, and never drinker), BMI (<18.5, 18.5–22.9, 23–24.9, 25–26.9,  $\geq$ 27), self-rated health (very good or good, fair, poor, and very poor; since no suicide death was observed in those with very good health, those with very good and good health were combined into one group), marital status (living with spouse, living without spouse), education (elementary school or less, middle school, and high school and more), monthly income (less than 1,000,000 Korean won [KRW], 1,000,000–1,990,000 KRW, and 2,000,000 KRW and more, where 1,300 KRW was about one US dollar as of June, 2001) and participant group (survey-only group and health exam group).

The sensitivity (proportion of true positives [suicides] correctly identified) and specificity (proportion of true negatives [non-suicides] correctly identified) at a cut-off score of 31 (or 30) were estimated and their 95% CI was calculated using the Wilson score method (*Newcombe, 1998*). The AUC was estimated using Proc Logistic (ROC statement) and averaged over 100 imputed data. A diagnostic test with an AUC value of 1.0 or 0.5 represents a perfect test or an uninformative test, respectively. Predictive values were not calculated because they are of little use to other populations with different suicide mortality.

The test for trend was done to demonstrate the dose–response relationships of the risk factors (every variable with three or more categories) to suicide by analyzing categories of a risk factor as ordinal variables. Additionally, analyses in all of the participants with follow-up until 2004, and analyses in survivors as of January 1, 2005 with follow-up until 2008, were done to evaluate whether the association of depressive symptoms and other risk factors with suicide, and the diagnostic characteristics of the BDI differ by follow-up period, and these calculations served as sensitivity (robustness) analyses. To test the non-response bias and potential harm of the screening for suicide, the suicide

mortality and age-adjusted hazard ratio (HR) of suicide of the participants (n = 10,238) were compared with those of the non-participants (n = 7,867).

All statistical analyses were performed using SAS version 9.4 (SAS Inc., Cary, North Carolina, USA). The *p*-value was calculated with two-sided tests. The proportional hazard assumption was tested using Martingale residuals and Schoenfeld residuals in complete data and imputed data analyses. No variables had evidence of a violation of the proportional hazard assumption except for a dummy variable of self-reported health (very good or good, only one suicide) in the complete data analysis.

## RESULTS

#### Characteristics of participants

The total follow-up person-years was 73,916. 41 men died by suicide (5.5 per 10,000 person-years, 95% CI [4.1–7.5]) during around 7.5 years of follow-up. The average (SD) age of the Korean veterans was 56.3 (3.5) years at enrollment. Participants with severe depressive symptoms were slightly younger than those with no-to-moderate symptoms, and they tended to be current smokers, past drinkers, slim, living without a spouse, unhealthy, and have little formal education and low earnings, compared to those with no-to-moderate symptoms (Table 1).

## Associations of depressive symptoms and other risk factors with suicide

In the unadjusted analysis, participants with severely depressive symptoms had a higher HR of suicide than those with no-to-moderately depressive symptoms, and increasing HR of suicide was associated with increasing depressive symptoms across the five categories of intensity (p for trend < 0.001) (Table 2, Table S1). Additionally, higher suicide risk correlated with a higher BDI score. After adjustment for various risk factors for suicide, the association of depressive symptoms with suicide (although somewhat weaker than before) remained strong.

Unadjusted HRs indicated that very poor health (p < 0.001), low education (p = 0.002), low household income (p = 0.03), past drinking (p = 0.07) and living without a spouse (p = 0.09) were associated with a higher risk of suicide, while obesity (BMI  $\ge 25$ ) with a lower risk of suicide (p = 0.07) (Table 3). In the unadjusted trend test across categories of risk factors, poorer self-rated health, lower education, and lower income were associated with a higher risk of suicide, while more obese subjects were less likely to die by suicide. After adjustment for depressive symptoms and other risk factors, very poor health, low education, and past drinking were still related to a higher HR of suicide, while BMI and marital status were not associated with suicide (Table 3, Table S6). In the adjusted trend test across categories of risk factors, only educational status remained statistically significantly associated with suicide (p = 0.03).

The associations of depressive symptoms with suicide were generally stronger during the earlier period of follow-up than during the later period (Table 4, Table S2). The associations of most of the other risk factors with suicide were also generally stronger (although none of them had a *p*-value below 0.05 due to the small number of suicide

#### Table 1 Characteristics of the Korean middle-aged male participants by depressive symptoms.

		Number (%)				
		Total	No to moderate symptoms	Severe symptoms	<i>p</i> value <sup>a</sup>	
Characteristics		<i>n</i> = 10,238	n = 7,470	n = 2,768		
Age at enrollment	y (SD)	56.3 (3.5)	56.5 (3.8)	56.2 (3.3)	0.007 <sup>b</sup>	
Smoking status <sup>c</sup>	Never smoker	3,027 (29.6)	2,333 (31.2)	694 (25.1)	< 0.001	
	Past smoker	2,480 (24.2)	1,783 (23.9)	698 (25.2)		
	Current smoker	4,730 (46.2)	3,354 (44.9)	1,377 (49.7)		
Drinking status	Never drinker	1,216 (11.9)	782 (10.5)	434 (15.7)		
	Past drinker	2,675 (26.1)	1,756 (23.5)	919 (33.2)	< 0.001	
	Current drinker	6,347 (62.0)	4,932 (66.0)	1,415 (51.1)		
BMI (kg/m <sup>2</sup> )	Below 18.5	323 (3.2)	160 (2.1)	163 (5.9)		
	18.5–22.9	4,118 (40.2)	2,843 (38.1)	1,275 (46.0)	< 0.001	
	23.0–24.9	2,992 (29.2)	2,296 (30.7)	696 (25.2)		
	25 or above	2,805 (27.4)	2,171 (29.1)	634 (22.9)		
Marital status <sup>c</sup>	Living with spouse	9,377 (91.6)	7,060 (94.5)	2,317 (83.7)	< 0.001	
	Living without spouse	861 (8.4)	410 (5.5)	450 (16.3)		
Self-rated health <sup>c</sup>	Very good or good	734 (7.2)	724 (9.7)	11 (0.4)		
	Fair	3,748 (36.6)	3,405 (45.6)	343 (12.4)	< 0.001	
	Poor	4,503 (44.0)	2,944 (39.4)	1,559 (56.3)		
	Very poor	1,253 (12.2)	398 (5.3)	856 (30.9)		
Educational status <sup>c</sup>	Elementary school or below	2,407 (23.5)	1,489 (19.9)	918 (33.2)		
	Middle school	2,782 (27.2)	1,934 (25.9)	848 (30.6)	< 0.001	
	High school or above	5,049 (49.3)	4,048 (54.2)	1,002 (36.2)		
Household income	Below 1,000,000	3,450 (33.7)	1,852 (24.8)	1,597 (57.7)		
per month <sup>c</sup>	1,000,000-1,990,000	4,693 (45.8)	3,707 (49.6)	987 (35.6)	< 0.001	
(Korean Won) <sup>d</sup>	2,000,000 or more	2,095 (20.5)	1,911 (25.6)	184 (6.7)		
Participant group	Survey-only group	8,233 (80.4)	5,881 (78.7)	2,352 (85.0)	< 0.001	
	Health exam group	2,005 (19.6)	1,589 (21.3)	416 (15.0)		
Depressive symptoms (1) <sup>e</sup>	No (0–13)	2,360 (23.0)	2,360 (31.6)	0 (0.0)		
(BDI score)	Mild (14–21)	2,554 (24.9)	2,554 (34.2)	0 (0.0)		
	Moderate (22–30)	2,556 (25.0)	2,556 (34.2)	0 (0.0)	< 0.001	
	Severe (31–39)	1,627 (15.9)	0 (0.0)	1,627 (58.8)		
	Extreme (40–63)	1,141 (11.1)	0 (0.0)	1,141 (41.2)		
Depressive symptoms (2) <sup>f</sup>	No (0–9)	1,307 (12.8)	1,307 (17.5)	0 (0.0)		
(BDI score)	Mild (10–16)	2,625 (25.6)	2,625 (35.1)	0 (0.0)		
	Moderate (17–29)	3,281 (32.0)	3,281 (43.9)	0 (0.0)	< 0.001	
	Severe (30–40)	2,015 (19.7)	257 (3.4)	1,758 (63.5)		
	Extreme (41–63)	1,010 (9.9)	0 (0.0)	1,010 (36.5)		

#### Notes.

BDI, Beck Depression Inventory; BMI, body mass index; SD, standard deviation.

<sup>a</sup> Combined *p*-value of chi-squared analysis over multiple imputed data.

<sup>b</sup> Combined p-value of analysis of variance over multiple imputed data.

<sup>c</sup> Sum of the number of participants from both groups may not equal the total due to rounding of averages over multiple imputed data.

<sup>d</sup> 1,300 Korean Won was about one US dollar as of June, 2001.

<sup>e</sup> Cut-off score based on quartiles and the last decile.

<sup>f</sup> Cut-off score based on previous research.

			Unadjusted analysis		Adjusted analysis <sup>b</sup>	
Categories of depressive symptoms (score)		No. of suicides <sup>c</sup>	<i>p</i> -value	HR (95% CI)	<i>p</i> -value	HR (95% CI)
Total BDI score (0–63)	Five-score increase	41	<0.001	1.4 (1.2–1.5)	0.01	1.2 (1.1-1.5)
2-group analysis <sup>d</sup>	No-to-moderate (0-30)	14		1.0 (Reference)		1.0 (Reference)
	Severe (31–63)	27	<0.001	5.4 (2.8–10.6)	0.004	3.4 (1.5–7.7)
5-group analysis <sup>d</sup>	No (0–13)	3		1.0 (Reference)		1.0 (Reference)
	Mild (14–21)	6	0.51	1.6 (0.4–6.8)	0.69	1.4 (0.3–6.1)
	Moderate (22-30)	5	0.70	1.4 (0.3–6.3)	0.94	0.9 (0.2–5.0)
	Severe (31–39)	12	0.009	5.4 (1.5–19.4)	0.12	3.3 (0.7–14.4)
	Extreme (40–63)	15	<0.001	10.2 (3.0–35.0)	0.05	4.6 (1.0-21.5)
	Trend test <sup>e</sup>	41	<0.001	1.9 (1.5–2.5)	0.009	1.6 (1.1–2.2)

#### Table 2 Hazard ratios of suicides by depressive symptoms in Korean middle-aged men (n = 10,238).<sup>a</sup>

#### Notes.

BDI, Beck Depression Inventory; CI, confidence interval; HR, hazard ratio.

<sup>a</sup> Hazard ratios were calculated using a Cox proportional hazards model over multiple imputed data.

<sup>b</sup> Adjusted for age at enrollment, smoking status, drinking status, body mass index, self-rated health, marital status, educational status, household monthly income and participant group.

<sup>c</sup> Number of suicides may not match those of the other groups of depressive symptoms due to rounding of averages over multiple imputed data.

<sup>d</sup> Cut-off score based on quartiles and the last decile.

<sup>e</sup> Five categories of Depressive symptoms were analyzed as ordinal variables.

deaths) during the earlier period than the later period, while educational status was more strongly associated with suicide during the later period than the earlier period (Table S3).

#### **Diagnostic properties of the Beck Depression Inventory**

For the unbinned total score and cut-off score of 31 or above, respectively, the AUC values were 0.71 and 0.69 during the follow-up, and they were higher during the first half of follow-up (2001–2004) than during the latter half (2005–2008) (Table 5, Fig. S2 and Table S4). The overall sensitivity and specificity of the 2-group test at a cut-off score of 31 or above on predicting suicide during 7.5 years of follow-up were 66% (95% CI [50–78]) and 73% (95% CI [72–74]), respectively. The sensitivity decreased over the follow-up period, while the specificity remained relatively unchanged. The AUC analyses showed that cut-off score of 31 or above had the highest AUC values across cut-off scores of BDI for predicting suicide (Table S5).

#### DISCUSSION

#### Depressive symptoms and suicide

Severe depressive symptoms were more strongly associated with suicide compared to no-to-moderate depressive symptoms in this study, after adjusting for other potential risk factors for suicide, although the association was weaker than that of depressive disorders examined in psychological autopsy studies (*Yoshimasu, Kiyohara & Miyashita, 2008*). The association of depressive symptoms with suicide was stronger during the first 3.5 years of follow-up than during the latter 4-year period in accordance with previous research (*Bramness et al., 2010; Sun et al., 2012*). This indicates that a cohort study of long follow-up with one baseline assessment and without considering the change in depressive symptoms

#### Table 3 Hazard ratios of suicides by risk factors in Korean middle-aged men (n = 10,238).<sup>a</sup>

			Unadjusted analysis		Adjusted analysis <sup>b</sup>	
Categories of risk factors		No. of suicides <sup>c</sup>	<i>p</i> -value	HR (95% CI)	<i>p</i> -value	HR (95% CI)
Age at enrollment	One year increase in age	41	0.24	0.9 (0.9–1.0)	0.27	0.9 (0.8–1.0)
Smoking status	Never smoker	9		1.0 (Reference)		1.0 (Reference)
	Past smoker	7	0.95	1.0 (0.4–2.6)	0.51	0.7 (0.3–2.0)
	Current smoker	25	0.15	1.8 (0.8–3.8)	0.47	1.3 (0.6–2.9)
	<i>p</i> for trend <sup>d</sup>	41	0.11	1.4 (0.9–2.0)	0.36	1.2 (0.8–1.8)
Drinking status	Never drinker	1		1.0 (Reference)		1.0 (Reference)
	Past drinker	16	0.07	8.4 (0.8-86.9)	0.05	8.7 (1.0–75.5)
	Current drinker	23	0.19	4.8 (0.5–49.1)	0.10	6.1 (0.7-53.0)
	<i>p</i> for trend <sup>d</sup>	41	0.80	1.1 (0.7–1.7)	0.29	1.3 (0.8–2.1)
Body mass index	Below 18.5	2	0.79	1.2 (0.3–5.2)	0.75	0.8 (0.2-3.4)
$(kg/m^2)$	18.5-22.9	22		1.0 (Reference)		1.0 (Reference)
	23.0–24.9	10	0.20	0.6 (0.3–1.3)	0.55	0.8 (0.4–1.7)
	25 or above	7	0.07	0.5 (0.2–1.1)	0.23	0.6 (0.2–1.4)
	<i>p</i> for trend <sup>d</sup>	41	0.04	0.7 (0.5–1.0)	0.29	0.8 (0.6–1.2)
Self-rated health	Very good or good	1	0.35	0.4 (0.1–2.9)	0.92	1.1 (0.1–10.3)
	Fair	9	0.33	0.7 (0.3–1.5)	0.71	1.2 (0.5–2.9)
	Poor	16		1.0 (Reference)		1.0 (Reference)
	Very poor	15	<0.001	3.6 (1.8–7.3)	0.02	2.4 (1.1–5.2)
	<i>p</i> for trend <sup>d</sup>	41	<0.001	2.4 (1.6–3.6)	0.13	1.5 (0.9–2.5)
Marital status	Living with spouse	35		1.0 (Reference)		1.0 (Reference)
	Living without spouse	6	0.09	2.1 (0.9–5.0)	0.76	1.2 (0.5–2.8)
Educational status	Elementary school or below	16	0.002	3.6 (1.6-8.2)	0.04	2.4 (1.0-5.6)
	Middle school	15	0.02	2.8 (1.2-6.3)	0.09	2.1 (0.9–4.9)
	High school or above	10		1.0 (Reference)		1.0 (Reference)
	<i>p</i> for trend <sup>d</sup>	41	<0.001	0.6 (0.4–0.8)	0.03	0.7 (0.5–1.0)
Household income	Below 1,000,000	21	0.03	3.7 (1.1–12.2)	0.76	1.2 (0.3–4.7)
per month	1,000,000–1,990,000	16	0.29	2.0 (0.6–6.8)	0.79	1.2 (0.3–4.4)
(Korean Won) <sup>e</sup>	2,000,000 or more	4		1.0 (Reference)		1.0 (Reference)
	<i>p</i> for trend <sup>d</sup>	41	0.01	0.5 (0.3–0.9)	0.80	0.9 (0.5–1.7)
Participant group	Health exam group	7	0.46	0.7 (0.3–1.7)	0.77	0.9 (0.4–2.1)
	Survey-only group	34		1.0 (Reference)		1.0 (Reference)

Notes.

BDI, Beck Depression Inventory; CI, confidence interval; HR, hazard ratio.

<sup>a</sup> Hazard ratios were calculated using a Cox proportional hazards model over multiple imputed data.

<sup>b</sup> Variables included in the Cox model were age at enrollment, smoking status, drinking status, body mass index, self-rated health, marital status, educational status, household monthly income, participant group, and depressive symptoms (5 categories based on quartiles and the last decile of the total BDI score).

<sup>c</sup> Sum of the number of suicides may not equal the total suicides (n = 41) due to rounding of averages over multiple imputed data.

<sup>d</sup> Categories of risk factors were analyzed as ordinal variables.

<sup>e</sup> 1,300 Korean Won was about one US dollar as of June, 2001.

during the follow-up, may underestimate the effects of depressive symptoms on suicide (*Bramness et al., 2010*). Additionally, in the current study, higher suicide risk was associated with a higher depressive symptom score in accordance with previous cohort studies in general-risk populations (*Bramness et al., 2010*; *Sun et al., 2012*). Overall, this prospective cohort study confirmed that depressive symptoms were a strong predictor of future suicide.

		From 2001 to 2004 ( <i>n</i> = 10,238)			From 2005 to 2008 ( <i>n</i> = 9,877)		
Categories of depressive symptoms (score)		No. of suicides <sup>b</sup>	<i>p</i> -value	HR (95% CI)	No. of suicides <sup>b</sup>	<i>p</i> -value	HR (95% CI)
Total BDI score (0–63)	Five-score increase	16	0.02	1.3 (1.1–1.8)	25	0.16	1.1 (1.0–1.4)
2-group analysis <sup>c</sup>	No-to-moderate (0–30)	4		1.0 (Reference)	10		1.0 (Reference)
	Severe (31–63)	12	0.02	5.2 (1.3–19.8)	15	0.06	2.7 (0.9–7.6)
5-group analysis <sup>c</sup>	No (0–13)	1		1.0 (Reference)	2		1.0 (Reference)
	Mild (14–21)	3	0.70	1.6 (0.1–17.6)	3	0.87	1.2 (0.2-8.4)
	Moderate (22-30)	1	0.99	0.0	4	0.80	1.3 (0.2–9.2)
	Severe (31–39)	4	0.33	3.3 (0.3–37.8)	8	0.21	3.3 (0.5–20.9)
	Extreme (40–63)	8	0.13	7.1 (0.6–88.3)	7	0.23	3.3 (0.5–23.8)
	Trend test <sup>d</sup>	16	0.03	1.9 (1.1–3.3)	25	0.09	1.4 (0.9–2.2)

#### Table 4 Adjusted hazard ratios of suicides by depressive symptoms according to follow-up period.<sup>a</sup>

Notes.

BDI, Beck Depression Inventory; CI, confidence interval; HR, hazard ratio.

<sup>a</sup> Hazard ratios were calculated using a Cox proportional hazards model over multiple imputed data, after adjustment for age at enrollment, smoking status, drinking status, body mass index, selfrated health, marital status, educational status, household monthly income and participant group.

<sup>b</sup> Number of suicides may not match those of the other groups of depressive symptoms due to rounding of averages over multiple imputed data.

<sup>c</sup> Cut-off score based on quartiles and the last decile.

<sup>d</sup> Five categories of depressive symptoms were analyzed as ordinal variables.

 Table 5 Diagnostic characteristics of Beck Depression Inventory by follow-up period in Korean middle-aged men.

Follow-up period	Characteristics	Cut-off of 31 or above <sup>a</sup> Rate (95% CI) <sup>b</sup>	Unbinned total score Rate (95% CI) <sup>b</sup>
From 2001 to 2008	Sensitivity <sup>c</sup> Specificity <sup>d</sup> AUC <sup>e</sup>	66 (50–78) 73 (72–74) 0.69 (0.62–0.77)	0.71 (0.63–0.80)
From 2001 to 2004	Sensitivity <sup>c</sup> Specificity <sup>d</sup> AUC <sup>e</sup>	75 (50–90) 73 (72–74) 0.74 (0.63–0.85)	0.75 (0.61–0.90)
From 2005 to 2008	Sensitivity <sup>c</sup> Specificity <sup>d</sup> AUC <sup>e</sup>	60 (41–76) 74 (73–75) 0.67 (0.57–0.77)	0.69 (0.58–0.80)

Notes.

AUC, area under the receiver operating characteristics curve; CI, confidence interval.

<sup>a</sup> Cutoff score based on third quartile.

<sup>b</sup> 95% CI was calculated using Wilson score method.

<sup>c</sup> Proportion of true positives (suicides) correctly identified.

<sup>d</sup> Proportion of true negatives (non-suicides) correctly identified.

<sup>e</sup> A diagnostic test with an AUC value of 1.0 or 0.5 represents a perfect test or an uninformative test, respectively.

#### **Diagnostic characteristics of Beck Depression Inventory**

To the best of our knowledge, this is the first study that has reported diagnostic characteristics for suicide death in a general-risk population. Although studies on suicidal ideation or attempt have been performed, attempters may share some properties with

suicides; however, they may be two different populations, especially among males (*Isometsa* & *Lonnqvist*, 1998; *Parra Uribe et al.*, 2013). The sensitivity and specificity at the cut-off score of 31 or above of the BDI in Korean middle-aged men for predicting future suicide, especially during the earlier period of follow-up, was comparable at least to those of other tools such as the Beck Hopelessness Scale (sensitivity 80%, specificity 42%) (*McMillan et al.*, 2007), Beck Suicide Intent Scale (sensitivity 77%, specificity 49%) (*Harriss & Hawton*, 2005), and a tool constructed from stepwise discriminant analysis of various instruments (sensitivity 56%, specificity 74%) (*Pokorny*, 1983) that were mainly applied to inpatients with psychiatric disorders or those who had done self-harm. They are also comparable to those of BDI at cut-off score of 23 or above in psychiatric outpatients (sensitivity 76%, specificity 62%) (*Beck et al.*, 1990) and those of BDI-II at cut-off score of 31 or above in prisoners (for attempted suicide or self-harm incident; sensitivity 80%, specificity 69%) (*Perry & Gilbody*, 2009). Although there is a room for improvement, the BDI may have an acceptable sensitivity and specificity as a suicide risk assessment tool.

It has been reported that male patients rarely initiated suicide-related discussions with their physician (*Vannoy & Robins, 2011*), while physicians were also reluctant to discuss suicide for various reasons including the fear of stimulating suicide in patients with depression by asking about it (*Crawford et al., 2011; Schulberg et al., 2004*). The suicide mortality of the participants in the present study (41 suicides, 5.5 suicides per 10,000 person-years) was not higher than that of the non-participants (34 suicides in 7,867 men, 6.0 (95% CI [4.3–8.4]) suicides per 10,000 person-years), and the age-adjusted hazard ratio of suicide (0.93, 95% CI [0.59–1.46]) also was not higher in the participants compared to non-participants. These results suggest that asking people about depressive symptoms using the BDI does not seem to be associated with a higher risk of suicide.

#### Other risk factors and suicide

Although physical health, marital status, drinking, and socioeconomic status have been associated with suicide in previous research, their association with suicide has rarely been examined after adjustment for depression (*Conwell, Duberstein & Caine, 2002*), especially in a prospective manner. Marital status was not associated with suicide in clinical patients with depression in a systematic review (*Hawton et al., 2013*), and it was also not associated with suicide after adjusting for other risk factors in a large psychological autopsy study in China (*Phillips et al., 2002*). Although BMI was inversely associated with suicide in large cohort studies, depressive symptoms were not accounted for in those studies (*Jee et al., 2011*; *Mukamal et al., 2010*). A review also concluded that much, if not all, of the association of physical health factors with suicide was mediated by affective disorders (*Conwell, Duberstein & Caine, 2002*).

In prospective studies, evidence linking regular alcohol consumption with suicide death is lacking (*Jee et al., 2011; Mukamal et al., 2007*), except for serious alcohol-related conditions such as alcohol use disorder and very heavy drinking (*Akechi et al., 2006; LeardMann et al., 2013*). Meanwhile, higher risks of suicide in past drinkers have also been identified in Japanese middle-aged men (depression was not adjusted for, though, in the Japanese

study) (*Akechi et al.*, 2006). As for socioeconomic status, despite a strong dose–response relationship of both educational status and household income with suicide in unadjusted analysis, the association of household income was substantially weakened after adjustment for self-rated health and depressive symptoms, because household income status was strongly associated with self-rated health and depressive symptoms in the current study.

Our findings suggest that very poor health, low education, and drinking status may be independently associated with suicide, while good or fair health, BMI, household income, and marital status may not be factors independently related factors to suicide. Instead, the associations of these non-independent factors with suicide may rather be mediated through depressive symptoms and other factors in Korean middle-aged men.

#### Limitations of the study

Despite of the strengths of the study, such as nearly complete prospective follow-up on suicide using national mortality data and the analysis of the diagnostic characteristics of a potential suicide risk assessment tool on suicide death in a general-risk population, this study has some limitations. First, our findings are based on 41 suicides, so the study may have lacked statistical power in some analyses. In particular, risk factors having a weak association with suicide may not be clearly identified in this study. Second, in cohort studies, nonresponse to the baseline survey may introduce bias. However, the suicide mortality of 7,867 (43%) non-respondents did not differ from that of participants, and item non-response to questionnaires was minimized with multiple imputation of 100 datasets. Third, the validity of the suicide death listed on death certificates was not examined separately. Since any misclassification on the suicide death could be most likely non-differential according to depressive symptoms, potential misclassifications would not substantially overestimate the hazard ratios.

It may be a limitation of its generalizability that this study's participants were Korean Vietnam War veterans. However, among US military personnel, mental health problems but not military-specific variables were independent risk factors for suicide (*LeardMann et al., 2013*). Additionally, the suicide mortality in Korean Vietnam War veterans did not differ from that from the male population of Korea (*Yi & Ohrr, 2011*). Therefore, we believe that performing the study with Vietnam War veterans did not substantially hinder its generalizability. Meanwhile, although psychological disorders such as depression have been universal risk factors among many cultural groups (*Sun et al., 2012; Zhang et al., 2004*), the strength of association and the association in itself of some risk factors with suicide may differ by ethnicity, culture, age, and gender (*Bjerkeset, Romundstad & Gunnell, 2008; Goldston et al., 2008; Milner et al., 2013; Sun et al., 2012; Zhang et al., 2004*). Therefore, some results in Korean middle-aged men may not be generalizable to other ethnic, cultural, or age groups, or to females.

#### **CONCLUSIONS**

Although the association was weaker than estimated from psychological autopsy studies, this study prospectively confirmed that depressive symptoms are a strong independent predictor of suicide. The sensitivity and specificity at the BDI cut-off score of 31 or above in Korean middle-aged men for predicting future suicide during 2001–2008 were 66% (95% CI [50–78]) and 73% (95% CI [72–74]), respectively. They were 75% (95% CI [50–90]) and 73% (95% CI [72–74]), respectively, for predicting future suicide during 2001–2004. BDI may have an acceptable sensitivity and specificity as a suicide risk assessment tool to help physicians and public health professionals identify individuals with depression and suicidal potential. This study also suggests that very poor self-rated health, drinking status and low education, but not marital status and BMI, may be independent predictors of suicide.

#### Abbreviations

AUC	area under the receiver operating characteristics curve
BDI	Beck Depression Inventory
BMI	body mass index
CI	confidence interval
HR	hazard ratio
KRW	Korean Won
ROC	the receiver operating characteristics

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## **Competing Interests**

The authors declare there are no competing interests.

#### **Author Contributions**

- Sang-Wook Yi conceived and designed the experiments, performed the experiments, analyzed the data, contributed reagents/materials/analysis tools, wrote the paper, prepared figures and/or tables, reviewed drafts of the paper.
- Jae-Seok Hong performed the experiments, reviewed drafts of the paper.

#### **Human Ethics**

The following information was supplied relating to ethical approvals (i.e., approving body and any reference numbers):

The veterans were provided with a written summary about the research and an invitation to the health exam by post. Return of the postal survey and voluntary participation of health exam were considered implied consent. This study was approved by the Institutional Review Board of Kwandong University (Approval No: KD-13-0105).

#### Supplemental Information

Supplemental information for this article can be found online at http://dx.doi.org/ 10.7717/peerj.1071#supplemental-information.

### REFERENCES

- Agerbo E. 2007. High income, employment, postgraduate education, and marriage: a suicidal cocktail among psychiatric patients. *Archives of General Psychiatry* **64**:1377–1384 DOI 10.1001/archpsyc.64.12.1377.
- Akechi T, Iwasaki M, Uchitomi Y, Tsugane S. 2006. Alcohol consumption and suicide among middle-aged men in Japan. *British Journal of Psychiatry* 188:231–236 DOI 10.1192/bjp.188.3.231.
- Al-Turkait FA, Ohaeri JU. 2010. Dimensional and hierarchical models of depression using the Beck Depression Inventory-II in an Arab college student sample. *BMC Psychiatry* 10:60 DOI 10.1186/1471-244X-10-60.
- Beck AT, Brown G, Berchick RJ, Stewart BL, Steer RA. 1990. Relationship between hopelessness and ultimate suicide: a replication with psychiatric outpatients. *American Journal of Psychiatry* 147:190–195 DOI 10.1176/ajp.147.11.1577-a.
- Beck AT, Steer RA. 1987. *Manual for the Beck Depression Inventory*. San Antonio: Psychological Corporation.
- Beck AT, Ward CH, Mendelson M, Mock J, Erbaugh J. 1961. An inventory for measuring depression. sion. Archives of General Psychiatry 4:561–571 DOI 10.1001/archpsyc.1961.01710120031004.
- Bjerkeset O, Romundstad P, Gunnell D. 2008. Gender differences in the association of mixed anxiety and depression with suicide. *British Journal of Psychiatry* 192:474–475 DOI 10.1192/bjp.bp.107.045203.
- Bramness JG, Walby FA, Hjellvik V, Selmer R, Tverdal A. 2010. Self-reported mental health and its gender differences as a predictor of suicide in the middle-aged. *American Journal of Epidemiology* 172:160–166 DOI 10.1093/aje/kwq091.
- Conwell Y, Duberstein PR, Caine ED. 2002. Risk factors for suicide in later life. *Biological Psychiatry* 52:193–204 DOI 10.1016/S0006-3223(02)01347-1.
- Crawford MJ, Thana L, Methuen C, Ghosh P, Stanley SV, Ross J, Gordon F, Blair G, Bajaj P.
   2011. Impact of screening for risk of suicide: randomised controlled trial. *British Journal of Psychiatry* 198:379–384 DOI 10.1192/bjp.bp.110.083592.
- Goldston DB, Molock SD, Whitbeck LB, Murakami JL, Zayas LH, Hall GC. 2008. Cultural considerations in adolescent suicide prevention and psychosocial treatment. *American Psychologist* 63:14–31 DOI 10.1037/0003-066X.63.1.14.

- Harriss L, Hawton K. 2005. Suicidal intent in deliberate self-harm and the risk of suicide: the predictive power of the Suicide Intent Scale. *Journal of Affective Disorders* 86:225–233 DOI 10.1016/j.jad.2005.02.009.
- Hawton K, Casanas ICC, Haw C, Saunders K. 2013. Risk factors for suicide in individuals with depression: a systematic review. *Journal of Affective Disorders* 147:17–28 DOI 10.1016/j.jad.2013.01.004.
- Hawton K, van Heeringen K. 2009. Suicide. *Lancet* 373:1372–1381 DOI 10.1016/S0140-6736(09)60372-X.
- Hjelmeland H, Dieserud G, Dyregrov K, Knizek BL, Leenaars AA. 2012. Psychological autopsy studies as diagnostic tools: are they methodologically flawed? *Death Studies* 36:605–626 DOI 10.1080/07481187.2011.584015.
- Isometsa ET, Lonnqvist JK. 1998. Suicide attempts preceding completed suicide. *British Journal of Psychiatry* 173:531–535 DOI 10.1192/bjp.173.6.531.
- Jee SH, Kivimaki M, Kang HC, Park IS, Samet JM, Batty GD. 2011. Cardiovascular disease risk factors in relation to suicide mortality in Asia: prospective cohort study of over one million Korean men and women. *European Heart Journal* 32:2773–2780 DOI 10.1093/eurheartj/ehr229.
- LeardMann CA, Powell TM, Smith TC, Bell MR, Smith B, Boyko EJ, Hooper TI, Gackstetter GD, Ghamsary M, Hoge CW. 2013. Risk factors associated with suicide in current and former US military personnel. *JAMA* 310:496–506 DOI 10.1001/jama.2013.65164.
- Masocco M, Pompili M, Vanacore N, Innamorati M, Lester D, Girardi P, Tatarelli R, Vichi M. 2010. Completed suicide and marital status according to the Italian region of origin. *Psychiatric Quarterly* 81:57–71 DOI 10.1007/s11126-009-9118-2.
- McMillan D, Gilbody S, Beresford E, Neilly L. 2007. Can we predict suicide and non-fatal self-harm with the Beck Hopelessness Scale? A meta-analysis. *Psychological Medicine* **37**:769–778 DOI 10.1017/S0033291706009664.
- Milner A, Hjelmeland H, Arensman E, De Leo D. 2013. Social-environmental factors and suicide mortality. *Sociology Mind* 3:137–148 DOI 10.4236/sm.2013.32021.
- Milner A, Page A, Lamontagne AD. 2014. Cause and effect in studies on unemployment, mental health and suicide: a meta-analytic and conceptual review. *Psychological Medicine* 44:909–917 DOI 10.1017/S0033291713001621.
- Mukamal KJ, Kawachi I, Miller M, Rimm EB. 2007. Drinking frequency and quantity and risk of suicide among men. *Social Psychiatry and Psychiatric Epidemiology* **42**:153–160 DOI 10.1007/s00127-006-0144-1.
- Mukamal KJ, Rimm EB, Kawachi I, O'Reilly EJ, Calle EE, Miller M. 2010. Body mass index and risk of suicide among one million US adults. *Epidemiology* 21:82–86 DOI 10.1097/EDE.0b013e3181c1fa2d.
- Newcombe RG. 1998. Interval estimation for the difference between independent proportions: comparison of eleven methods. *Statistics in Medicine* 17:873–890 DOI 10.1002/(SICI)1097-0258(19980430)17:8 < 873::AID-SIM779 > 3.0.CO;2-I.
- Parra Uribe I, Blasco-Fontecilla H, Garcia-Pares G, Giro Batalla M, Llorens Capdevila M, Cebria Meca A, de Leon-Martinez V, Perez-Sola V, Palao Vidal DJ. 2013. Attempted and completed suicide: not what we expected? *Journal of Affective Disorders* 150:840–846 DOI 10.1016/j.jad.2013.03.013.
- **Perry AE, Gilbody S. 2009.** Detecting and predicting self-harm behaviour in prisoners: a prospective psychometric analysis of three instruments. *Social Psychiatry and Psychiatric Epidemiology* **44**:853–861 DOI 10.1007/s00127-009-0007-7.

- Phillips MR, Yang G, Zhang Y, Wang L, Ji H, Zhou M. 2002. Risk factors for suicide in China: a national case-control psychological autopsy study. *Lancet* 360:1728–1736 DOI 10.1016/S0140-6736(02)11681-3.
- **Pokorny AD. 1983.** Prediction of suicide in psychiatric patients. Report of a prospective study. *Archives of General Psychiatry* **40**:249–257 DOI 10.1001/archpsyc.1983.01790030019002.
- Rhee M, Lee Y, Park S, Sohn C, Chung Y, Hong S, Lee B, Chang P, Yoon A. 1995. A standardization study of Beck Depression Inventory I-Korean version (K-BDI): reliability and factor analysis. *Korean Journal of Psychopathology* **4**:77–95.
- Schulberg HC, Bruce ML, Lee PW, Williams Jr JW, Dietrich AJ. 2004. Preventing suicide in primary care patients: the primary care physician's role. *General Hospital Psychiatry* 26:337–345 DOI 10.1016/j.genhosppsych.2004.06.007.
- Smarr KL, Keefer AL. 2011. Measures of depression and depressive symptoms: Beck Depression Inventory-II (BDI-II), Center for Epidemiologic Studies Depression Scale (CES-D), Geriatric Depression Scale (GDS), Hospital Anxiety and Depression Scale (HADS), and Patient Health Questionnaire-9 (PHQ-9). Arthritis Care & Research 63(Suppl 11):S454–S466 DOI 10.1002/acr.20556.
- Sun WJ, Xu L, Chan WM, Lam TH, Schooling CM. 2012. Depressive symptoms and suicide in 56,000 older Chinese: a Hong Kong cohort study. *Social Psychiatry and Psychiatric Epidemiology* 47:505–514 DOI 10.1007/s00127-011-0362-z.
- Vannoy SD, Robins LS. 2011. Suicide-related discussions with depressed primary care patients in the USA: gender and quality gaps. A mixed methods analysis. *BMJ Open* 1:e000198 DOI 10.1136/bmjopen-2011-000198.
- White IR, Royston P, Wood AM. 2011. Multiple imputation using chained equations: issues and guidance for practice. *Statistics in Medicine* **30**:377–399 DOI 10.1002/sim.4067.
- **World Health Organization. 2014.** *Preventing suicide: a global imperative.* Geneva: World Health Organization.
- Yi SW, Ohrr HC. 2011. Mortality patterns among Korean Vietnam Veterans. *Korean Journal of Occupational and Environmental Medicine* 23:279–286.
- Yi SW, Ryu SY, Ohrr H, Hong JS. 2014. Agent Orange exposure and risk of death in Korean Vietnam veterans: Korean Veterans Health Study. *International Journal of Epidemiology* 43:1825–1834 DOI 10.1093/ije/dyu183.
- Yoshimasu K, Kiyohara C, Miyashita K, Stress Research Group of the Japanese Society for Hygiene. 2008. Suicidal risk factors and completed suicide: meta-analyses based on psychological autopsy studies. *Environmental Health and Preventive Medicine* 13:243–256 DOI 10.1007/s12199-008-0037-x.
- Zhang J, Conwell Y, Zhou L, Jiang C. 2004. Culture, risk factors and suicide in rural China: a psychological autopsy case control study. *Acta Psychiatrica Scandinavica* 110:430–437 DOI 10.1111/j.1600-0447.2004.00388.x.