

Article

Association between Psychological Factors and Evacuation Status and the Incidence of Cardiovascular Diseases after the Great East Japan Earthquake: A Prospective Study of the Fukushima Health Management Survey

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Abstract: Evidence regarding the effect of psychological factors and evacuation on cardiovascular disease occurrence after large-scale disasters is limited. This prospective study followed up a total of 37,810 Japanese men and women aged 30–89 years from the Fukushima Prefecture with no history of stroke or heart disease at baseline (2012), until 2017. This period included 3000 cardiovascular events recorded through questionnaires and death certificates. The participants' psychological distress, trauma reaction, and evacuation status were defined, and divided into four groups based on combinations of psychological factors and evacuation status. We calculated the hazard ratios and 95% confidence intervals for only psychological, only evacuation, or both of them compared with neither using Cox proportional hazard models. Psychological factors along with evacuation resulted in approximately 5% to 25% higher magnitude of stroke and heart disease risk than psychological factors only among men. Compared to neither, the multivariable hazard ratios of those with both psychological distress and evacuation were 1.75 for stroke and 1.49 for heart disease, and those of both trauma reaction and evacuation were 2.01 and 1.57, respectively, among men. Evacuation combined with psychological factors increased the risk of stroke and heart disease risks especially in men after the Great East Japan Earthquake.

Keywords: Great East Japan Earthquake; disaster; cardiovascular disease; psychological factors; evacuation; prospective study

1. Introduction

The Great East Japan Earthquake on March 11, 2011 registered 9.0 on the Richter scale and caused a tsunami, resulting in a nuclear disaster in Fukushima Prefecture. Consequently, the prefectural government established evacuation zones in Fukushima, and more than 160,000 residents were forced to evacuate. As of January 14, 2020, more than 48,000 residents are still evacuated and bear psychological burdens and concerns regarding the lingering radiation risks. The local government therefore launched the Fukushima Health Management Survey (FHMS) to investigate and monitor the evacuees' health condition [1]. To date, numerous studies have examined the relationship between disasters and cardiovascular disease (CVD) in individuals who experience such events [2–6]. Earlier studies on the Great East Japan Earthquake have revealed that the incidence of CVD, including out-of-hospital cardiac arrest, tachyarrhythmias, heart failure, acute coronary syndrome, and stroke, significantly increased following the disaster [7–9].

There is a complex association between disaster occurrence and CVD risk, and the available research indicates numerous risk factors, such as lifestyle-related, social, and psychological factors [10–12]. A number of studies have reported that the disaster was followed by cases of serious post-traumatic stress disorder (PTSD), which in turn induced CVD in certain individuals affected by the disaster [13,14]. According to evidence from a few studies, various modifiable risk factors for CVD in addition to psychological factors have been identified, one of which is evacuation status [15–17]. After a disaster, the resulting traumatic events and evacuation status affect individuals' lifestyle and psychological state [18,19]. Earlier studies on the Great East Japan Earthquake have shown that evacuation status increases the incidence of hypertension, obesity, metabolic syndrome, diabetes, hyperlipidemia, liver dysfunction, chronic kidney disease, and polycythemia, all of which are CVD risk factors [20–27]. Although a number of studies have focused on the relationship between psychological factors and CVD risk or between evacuation status and CVD risk, they have not clarified how evacuation status affects the association between psychological factors and CVD occurrence.

We therefore hypothesized that, since the occurrence of the Great East Japan Earthquake, psychological factors have become risk factors for CVD and that evacuation status might increase the effect of these factors. To our knowledge, no previous prospective study has examined the combined effect of psychological factors and evacuation status on CVD after the occurrence of large-scale disasters among worldwide populations. This large cohort study therefore prospectively examined the combined effects of psychological factors and evacuation status on CVD occurrence among male and female Japanese residents who experienced the earthquake in the Fukushima prefecture.

2. Materials and Methods

2.1. Participants

To monitor the health status of Fukushima evacuees after the earthquake and to provide them with appropriate care, the Fukushima Medical University has been conducting an annual self-administered survey titled the "Mental Health and Lifestyle Survey [1]." One of its baseline surveys was conducted in January 2012 to investigate CVD onset, psychological factors, evacuation status, and lifestyle-related factors. Figure 1 provides a flow chart of the participant selection for the present study. In the survey, 64,668 Japanese individuals (28,687 men, 35,981 women) aged 30–89 years participated from 13 municipalities in the Fukushima Prefecture. Follow-up surveys assessing CVD onset were subsequently conducted once a year from baseline, and questionnaires were mailed to the participants annually. The present study excluded 2,633 men and 4,501 women whose history of psychological factors or evacuation status in 2012 was missing or unidentifiable, as well as 385 men and 596 women whose CVDs history in 2012 was missing or unidentifiable. The study also excluded 4,111 men and

3,394 women who already had a history of CVDs in 2012. The study further excluded 5,202 men and 6,036 women who never responded to the follow-up surveys. The final analysis therefore included 37,810 participants (16,356 men, 21,454 women).

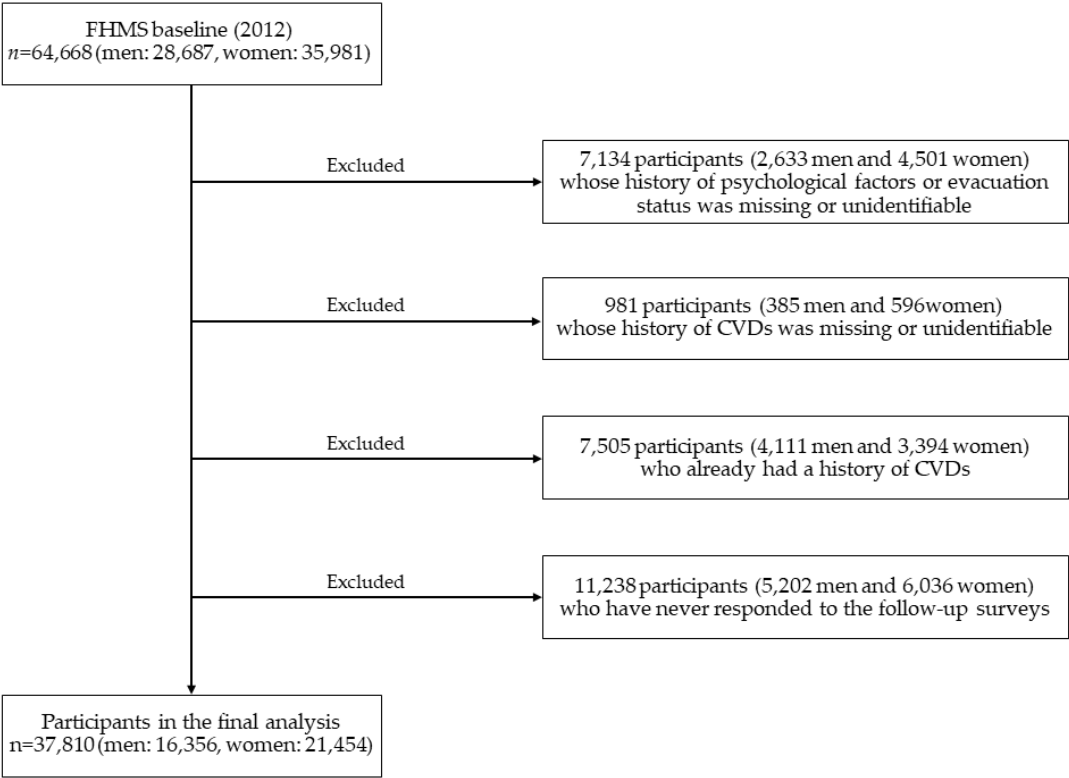


Figure 1. Flow chart of participant selection for the present study of association between psychological factors and evacuation status and the incidence of cardiovascular diseases after the Great East Japan Earthquake

The study was conducted based on the provisions of the Declaration of Helsinki, and the study protocol was approved by the Ethics Review Committees of Okayama University (No. 1803-022) and Fukushima Medical University (No. 1316, No. 2148, No. 2020-047). Submission of the self-administered questionnaires was considered as consent given by the participants for participating in the study.

2.2. Measurements

2.2.1. Onset and mortality of cardiovascular diseases

The study considered both onset and death as cardiovascular incidents. To measure the onset of CVDs (stroke or heart disease), the participants were presented once a year with the question, “Have you ever been diagnosed by a doctor with the following diseases?” and were asked to circle either “yes” or “no” for any applicable diseases [15,28–30]. The mortality data were forwarded to the public health departments of the respective areas before being centralized at the Ministry of Health, Labour and Welfare, and the underlying causes of death were coded based on the 10th revision of the International Classification for Diseases (ICD-10). The primary endpoints for the current analysis were death from stroke (ICD-10 codes I60–I69), heart disease (ICD-10 codes I20–I25 and I30–I52), and total CVD (stroke and heart disease).

2.2.2. Psychological factors and evacuation status

To assess the participants’ mental health status, we used the six-item Kessler Psychological Distress Scale (K6) and the Post-Traumatic Stress Disorder (PTSD) Checklist—Stressor-Specific Version

(PCL-S). In particular, the K6 was employed to measure psychological distress and screen individuals for nonspecific serious mental illness [31]. The scale included questions on whether the participants had experienced any of the following six symptoms during the past 30 days: "feeling so sad that nothing could cheer you up," "feeling nervous," "feeling hopeless," "feeling restless or fidgety," "feeling everything was an effort," and "feeling worthless." Each question was scored on a five-point Likert-type scale, with values ranging from 0 to 4. Scores ranged from 0 to 24, and higher scores indicated lower mental health status. The study validated and employed the Japanese version of K6 [32,33]. We defined psychological distress as scores ≥ 13 .

The PCL-S was employed to assess the reaction to trauma to identify the participants' current trauma-related symptoms [34]. The scale is a 17-item self-administered measure that detects PTSD, where each item is scored from 1 to 5 according to the responses "not at all," "a little bit," "moderately," "quite a bit," or "extremely," respectively. Total scores range from 17 to 85, with higher scores indicating greater reaction to trauma. The study validated and employed the Japanese version of PCL-S [29,30]. We classified the participants with PCL-S scores ≥ 44 as having trauma reaction.

Evacuation status was assessed based on the living conditions category. Participants were asked to select an answer from six options on their current living conditions: "evacuation shelter," "temporary housing," "rental housing or apartment," "a relative's home," "their own home," or "other." We defined those participants as evacuees who were currently living in or had lived in either an "evacuation shelter" or "temporary housing."

2.2.3. Lifestyle behaviors and social factors

Lifestyle behaviors and social factors were employed as adjusted variables for the association between psychological factors or evacuation status and cardiovascular incidents. Lifestyle behaviors included the participants' smoking status, alcohol consumption, physical activity, and sleep quality. We assessed the participants' smoking status using the question "Do you smoke (anything other than cigars and pipes)?" with the following options: "non-smoker", "ex-smoker", and "current smoker". Those who selected "current smoker" were considered current smokers. For the alcohol consumption category, individuals were asked, "Do you consume alcohol?", and the options were "less than once per month," "ex-drinker," and "once or more per month". Those who selected "once or more per month" were considered to have an alcohol consumption of once or more per month. The participants' physical activity level was assessed by the question "Do you exercise regularly?" with the following options: " \geq daily," "2-4 times/week," "weekly" and "never". Those who selected "weekly" were considered to have a physical activity frequency of once or more a week. For the sleep quality category, the question was "Are you satisfied with the quality of sleep for the past month (regardless of the length of sleep)?", and the options were "satisfied," "slightly dissatisfied," "very dissatisfied," and "extremely dissatisfied." Those who selected "slightly dissatisfied," "very dissatisfied," or "extremely dissatisfied" were considered unsatisfied with sleep.

The study considered job loss, loss of family members, and a high perception of radiation risks to be social factors. The "job loss" and "loss of family members" categories included the following "yes" or "no" questions: "Did you become unemployed?" and "Did you lose loved ones in this earthquake?" A "yes" response to these questions was considered to be confirmation of job loss and loss of family members. Furthermore, for the "high perception of radiation risks" category, participants answered the following multiple-choice questions: "How likely do you think acute health problems (e.g., death within a month) will occur from the current radiation exposure?," "How likely do you think health problems (e.g., cancer) will occur in the coming years due to the current radiation exposure?," and "To what extent do you think the current radiation exposure will affect future generations (children, grandchildren, etc.)?" The participants who answered "3" or "4" (where 1 corresponded to a low likelihood and 4 to a high likelihood) to any of these questions were considered to have a high perception of radiation risks.

2.3. Statistical Analysis

The age-adjusted mean and prevalence of baseline variables of interest were compared between participants according to evacuation status, using analysis of covariance and logistic regression models

[35]. Furthermore, *p* for differences were calculated between evacuees and non-evacuees. The number of cohorts was sufficient for the current analysis, and the risk of stroke was more significant among men; hence, the analyses in this study were stratified by gender.

The person-years of follow-up were calculated from the date of the response to the baseline questionnaire to the attainment of one of the following three possible endpoints: (1) a CVD event incidence (including death); (2) relocation from the study area; or (3) date of last response to the self-administered survey.

The participants were divided into four groups: participants with (1) neither evacuation status nor psychological factors (neither), (2) only psychological factors (only psychological), (3) only evacuation status (only evacuation), and (4) those with both psychological factors and evacuation status (both). The hazard ratios (HRs) and 95% confidence intervals (CIs) of cardiovascular incidence for the groups of only psychological, only evacuation, or both compared to neither were estimated using Cox proportional hazards models according to gender or evacuation status; subsequently, the relationship between each risk factors such as psychological distress or trauma reaction, and CVDs was examined. Similarly, the relationship between evacuation status and CVDs was evaluated. Furthermore, we investigate in detail the impact of gender or evacuation status-specific effect modifications on the association between psychological factors, such as psychological distress or trauma reaction, and CVDs. The *p*-values for interactions by gender or evacuation status were tested using the cross-product terms of gender and psychological distress or trauma reaction, or evacuation status and psychological distress or trauma reaction.

The adjustment variables included gender (dichotomous), age (continuous), smoking status (dichotomous), alcohol consumption (dichotomous), physical activity (dichotomous), sleep quality (dichotomous), and job loss (dichotomous). We used SAS Version 9.4 (SAS Institute, Inc., Cary, NC) for the statistical analysis. In this study, all statistical tests were two-tailed, and values of *p* < 0.05 were considered significant.

3. Results

During the 3.7-year mean follow-up period, the onset of all CVDs was reported for 2,829 (1,511 men and 1,318 women), including 626 stroke onset cases (361 men and 265 women); 2,203 heart disease onset cases (1,150 men and 1,053 women); and the death of 171 participants from all CVDs (86 men and 85 women), including 72 stroke (26 men and 46 women) and 99 heart disease (60 men and 39 women) death cases.

3.1. Cardiovascular risk factors of participants at baseline

Table 1 depicts the age-adjusted mean or prevalence of cardiovascular risk factors for the participants at baseline according to their evacuation status. Compared to non-evacuees, the average age of evacuees was lower, and evacuees had a higher prevalence of psychological distress, trauma reaction, current smoker, unsatisfied with sleep, job loss, loss of family members, and a high perception of radiation risks both in men and women.

Table 1. Age-adjusted mean or prevalence of cardiovascular risk factors for the participants at baseline according to their evacuation status.

	Evacuee	Non-evacuee	<i>p</i> for difference
Men			
Number at risk	5,990	10,336	
Age (years)	57.6	58.4	<0.001
Psychological distress (%)	13.6	9.2	<0.001
Trauma reaction (%)	21.1	14.4	<0.001
Current smoker (%)	35.6	33.2	0.03
Alcohol consumption ≥ once/month (%)	68.6	71.5	<0.001
Physical activity ≥ once/month (%)	47.8	47.5	0.38

Unsatisfied with sleep (%)	64.9	56.5	<0.001
Job loss (%)	28.0	13.9	<0.001
Loss of family members (%)	23.6	16.9	<0.001
High perception of radiation risks (%)	61.4	56.4	<0.001
Women			
Number at risk	8,172	13,282	
Age (years)	55.7	56.5	<0.001
Psychological distress (%)	18.5	13.9	<0.001
Trauma reaction (%)	27.1	20.5	<0.001
Current smoker (%)	10.7	9.3	0.02
Alcohol consumption ≥ once/month (%)	30.6	71.5	0.38
Physical activity ≥ once/month (%)	47.8	47.5	0.002
Unsatisfied with sleep (%)	64.9	56.5	<0.001
Job loss (%)	28.0	13.9	<0.001
Loss of family members (%)	23.6	16.9	<0.001
High perception of radiation risks (%)	61.4	56.4	<0.001

3.2. Psychological factors or evacuation status on cardiovascular diseases

Table 2 depicts the psychological factors or evacuation status-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to gender. For psychological distress, both men and women had an increased risk of stroke and heart disease. The respective multivariable HRs (95% CIs) were 1.53 (1.22–1.92) and 1.40 (1.22–1.60) for men and 1.39 (1.12–1.73) and 1.40 (1.24–1.57) for women. Similarly, for trauma reaction, both men and women showed an increased risk of stroke and heart disease. Respective multivariable HRs were 1.78 (1.49–2.14) and 1.43 (1.28–1.59) for men and 1.41 (1.17–1.70) and 1.35 (1.22–1.50) for women. For evacuation status, men showed significant or borderline significant increased risks for stroke and heart disease. The respective multivariable HRs were 1.17 (0.98–1.40) and 1.11 (1.00–1.23). Women showed no increase in risk of stroke or heart disease based on their evacuation status. There were statistically significant interactions based on gender between evacuation status and stroke ($p = 0.04$).

Table 2. Psychological factors or evacuation status-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to gender.

	Men	Women	<i>p</i> for interaction ^a
Person-years	59,087	79,350	
Psychological distress			
Total CVD (<i>n</i>)	301	449	
Incidence rate/1000 person-years	5.09	5.66	
Age-adjusted HR	1.48 (1.31–1.67)	1.48 (1.33–1.64)	0.93
Multivariable-adjusted HR ^b	1.34 (1.18–1.52)	1.39 (1.25–1.55)	0.90
Stroke (<i>n</i>)	92	111	
Incidence rate/1000 person-years	1.56	1.40	
Age-adjusted HR	1.66 (1.33–2.07)	1.48 (1.20–1.83)	0.45
Multivariable-adjusted HR	1.53 (1.22–1.92)	1.39 (1.12–1.73)	0.49
Heart disease (<i>n</i>)	262	381	
Incidence rate/1000 person-years	4.43	4.80	
Age-adjusted HR	1.56 (1.37–1.77)	1.49 (1.33–1.67)	0.59
Multivariable-adjusted HR	1.40 (1.22–1.60)	1.40 (1.24–1.57)	0.73
Trauma reaction			
Total CVD (<i>n</i>)	523	662	
Incidence rate/1000 person-years	8.85	8.34	
Age-adjusted HR	1.54 (1.40–1.70)	1.43 (1.30–1.57)	0.37

Multivariable-adjusted HR	1.42 (1.29–1.52)	1.35 (1.23–1.49)	0.41
Stroke (n)	174	170	
Incidence rate/1000 person-years	2.94	2.14	
Age-adjusted HR	1.86 (1.56–2.22)	1.49 (1.24–1.79)	0.09
Multivariable-adjusted HR	1.78 (1.49–2.14)	1.41 (1.17–1.70)	0.08
Heart disease (n)	437	559	
Incidence rate/1000 person-years	7.40	7.04	
Age-adjusted HR	1.56 (1.40–1.73)	1.44 (1.30–1.59)	0.39
Multivariable-adjusted HR	1.43 (1.28–1.59)	1.35 (1.22–1.50)	0.45
Evacuation status			
Total CVD (n)	871	819	
Incidence rate/1000 person-years	14.74	10.32	
Age-adjusted HR	1.18 (1.09–1.29)	1.07 (0.98–1.17)	0.10
Multivariable-adjusted HR	1.13 (1.02–1.24)	1.03 (0.94–1.14)	0.13
Stroke (n)	247	182	
Incidence rate/1000 person-years	4.18	2.29	
Age-adjusted HR	1.21 (1.03–1.42)	0.93 (0.77–1.11)	0.03
Multivariable-adjusted HR	1.17 (0.98–1.40)	0.90 (0.74–1.10)	0.04
Heart disease (n)	719	698	
Incidence rate/1000 person-years	12.17	8.80	
Age-adjusted HR	1.17 (1.06–1.28)	1.10 (1.00–1.21)	0.29
Multivariable-adjusted HR	1.11 (1.00–1.23)	1.04 (0.93–1.16)	0.35

Abbreviations: CVD, cardiovascular disease; HR, hazards ratio; CI, confidence interval. Notes: Values in parentheses indicate 95% confidence intervals. ^a *p* for interaction was calculated for the cross-product terms of gender and psychological distress, trauma reaction or evacuation status on CVDs. ^b Adjusted for age, smoking status, alcohol consumption, physical activity, sleep quality, and job loss.

3.3. Combination of psychological factors and evacuation status on cardiovascular diseases

Table 3 depicts the gender-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to the combination of psychological factors and evacuation status. Compared to those with neither psychological distress nor evacuation status, those with both had increased risks of stroke and heart disease, and the increased magnitude due to evacuation was approximately 0.21 for stroke and 0.05 for heart disease in men. The respective multivariable HRs (95% CIs) were 1.75 (1.26–2.44) for stroke and 1.49 (1.22–1.82) for heart disease. Compared to those with neither trauma reaction nor evacuation status, those with both had increased risks of stroke and heart disease, and the increased magnitude due to evacuation was approximately 0.25 for stroke and about 0.19 for heart disease in men. The respective multivariable HRs were 2.01 (1.54–2.61) for stroke and 1.57 (1.34–1.84) for heart disease. Evacuation status in addition to psychological factors did not lead to an additional impact on CVDs in women.

Table 3. Gender-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to the combination of psychological factors and evacuation status.

	Neither	Only psychological	Only evacuation	Both
Men				
Person-years	31,101	6,218	16,702	5,067
Psychological distress				
Total CVD (n)	1,162	156	726	145
Age-adjusted HR	1.00	1.50 (1.27–1.77)	1.17 (1.07–1.29)	1.65 (1.39–1.96)
Multivariable-adjusted HR ^a	1.00	1.37 (1.15–1.62)	1.13 (1.02–1.25)	1.46 (1.22–1.76)
Stroke (n)	328	47	202	45
Age-adjusted HR	1.00	1.65 (1.22–2.25)	1.18 (0.99–1.41)	1.89 (1.38–2.58)

Multivariable-adjusted HR	1.00	1.54 (1.13–2.10)	1.17 (0.96–1.41)	1.75 (1.26–2.44)
Heart disease (n)	960	138	595	124
Age-adjusted HR	1.00	1.60 (1.34–1.91)	1.16 (1.05–1.28)	1.70 (1.41–2.05)
Multivariable-adjusted HR	1.00	1.44 (1.21–1.73)	1.11 (0.99–1.24)	1.49 (1.22–1.82)
Trauma reaction				
Total CVD (n)	1,057	261	609	262
Age-adjusted HR	1.00	1.48 (1.30–1.71)	1.13 (1.02–1.25)	1.75 (1.53–2.00)
Multivariable-adjusted HR	1.00	1.38 (1.21–1.59)	1.09 (0.98–1.22)	1.58 (1.37–1.84)
Stroke (n)	287	88	161	86
Age-adjusted HR	1.00	1.83 (1.44–2.32)	1.14 (0.94–1.38)	2.08 (1.64–2.65)
Multivariable-adjusted HR	1.00	1.76 (1.38–2.24)	1.13 (0.92–1.39)	2.01 (1.54–2.61)
Heart disease (n)	880	218	500	219
Age-adjusted HR	1.00	1.49 (1.28–1.73)	1.11 (0.99–1.24)	1.76 (1.52–2.04)
Multivariable-adjusted HR	1.00	1.38 (1.19–1.60)	1.07 (0.95–1.20)	1.57 (1.34–1.84)
Women				
Person-years	37,141	11,498	21,371	9,339
Psychological distress				
Total CVD (n)	1,002	257	627	192
Age-adjusted HR	1.00	1.53 (1.34–1.76)	1.08 (0.98–1.19)	1.50 (1.29–1.75)
Multivariable-adjusted HR	1.00	1.45 (1.26–1.66)	1.05 (0.94–1.17)	1.38 (1.18–1.63)
Stroke (n)	265	64	135	47
Age-adjusted HR	1.00	1.44 (1.09–1.89)	0.89 (0.73–1.10)	1.41 (1.03–1.92)
Multivariable-adjusted HR	1.00	1.34 (1.01–1.77)	0.87 (0.70–1.09)	1.29 (0.93–1.79)
Heart disease (n)	832	218	522	163
Age-adjusted HR	1.00	1.56 (1.35–1.82)	1.11 (0.99–1.23)	1.53 (1.29–1.81)
Multivariable-adjusted HR	1.00	1.47 (1.26–1.71)	1.06 (0.94–1.19)	1.38 (1.15–1.65)
Trauma reaction				
Total CVD (n)	880	379	536	283
Age-adjusted HR	1.00	1.55 (1.37–1.75)	1.11 (1.00–1.24)	1.41 (1.23–1.61)
Multivariable-adjusted HR	1.00	1.47 (1.30–1.66)	1.08 (0.97–1.22)	1.30 (1.13–1.50)
Stroke (n)	225	104	116	283
Age-adjusted HR	1.00	1.63 (1.30–2.06)	0.97 (0.77–1.21)	1.27 (0.97–1.67)
Multivariable-adjusted HR	1.00	1.55 (1.22–1.96)	0.95 (0.75–1.20)	1.17 (0.88–1.57)
Heart disease (n)	735	315	454	244
Age-adjusted HR	1.00	1.54 (1.35–1.76)	1.13 (1.00–1.27)	1.46 (1.26–1.69)
Multivariable-adjusted HR	1.00	1.45 (1.27–1.66)	1.08 (0.95–1.23)	1.32 (1.13–1.54)

253 Abbreviations: CVD, cardiovascular disease; HR, hazards ratio; CI, confidence interval. Note: ^a Adjusted for
254 age, smoking status, alcohol consumption, physical activity, sleep quality, and job loss.

255 Supplementary Table S1 depicts the gender-specific age-adjusted and multivariable HRs (95% CIs)
256 of CVDs according to evacuation status. Men with psychological distress showed an increased risk
257 of both stroke and heart disease, irrespective of evacuation status. The respective multivariable HRs
258 (95% CIs) were 1.49 (1.07–2.08) for evacuees and 1.54 (1.13–2.11) for non-evacuees for stroke, and
259 1.39 (1.14–1.69) vs 1.41 (1.18–1.70) for heart disease. Similarly, men with trauma reaction showed an
260 increased risk, and respective multivariable HRs of 1.79 (1.36–2.35) vs 1.75 (1.37–2.24) for stroke,
261 and 1.52 (1.29–1.79) vs 1.35 (1.16–1.57) for heart disease. Women with psychological distress
262 showed a significant or borderline significant increased risk of both stroke and heart disease,
263 irrespective of evacuation status. The respective multivariable HRs (95% CIs) were 1.50 (1.07–2.11)
264 for evacuees and 1.32 (1.00–1.74) for non-evacuees for stroke, and 1.30 (1.09–1.56) vs 1.47 (1.26–1.71)
265 for heart disease. Women with a trauma reaction showed an increased risk of stroke for non-
266 evacuees and heart disease irrespective of evacuation status. The respective multivariable HRs (95%
267 CIs) were 1.53 (1.21–1.95) for non-evacuees for stroke, and 1.22 (1.04–1.43) vs 1.46 (1.27–1.67) for
268 heart disease.

269 **4. Discussion**

270 Using data from a large-scale prospective study among Japanese men and women aged 30–89
271 years in Fukushima after the Great East Japan Earthquake, we observed an association between
272 psychological factors and evacuation status, and CVD risk among individuals who experienced the
273 disaster. Psychological factors along with evacuation status resulted in approximately 5% to 25%
274 higher magnitude of stroke and heart disease risk in men than psychological factors alone.

275 The increased CVD risk caused by psychological factors found in our study is consistent with
276 the findings of earlier prospective cohort studies on individuals' experiences following traumatic
277 events such as disasters [12–14]. After the 2001 World Trade Center Disaster, a prospective cohort
278 study was conducted in New York State, in which 46,346 participants were followed up for a mean
279 period of 6.5 years per person. The adjustment HRs of patients with PTSD were approximately 1.5
280 times higher in men for cerebrovascular disease and 1.3 times higher in women for heart disease than
281 for their counterparts without PTSD [36]. There have been similar reports on the relationship between
282 psychological factors and CVDs following earthquake disasters as well. In a prospective cohort study
283 conducted in China following the 2008 Sichuan earthquake, 404 participants were followed up for 2.0
284 years. This study found that the odds of participants having depressive symptoms caused by
285 earthquake-related loss was approximately 1.6 times higher than in those who did not experience
286 earthquake-related loss [37]. Furthermore, several studies have examined evacuation status
287 following traumatic events and reported similar relationships [15–17]. In the FHMS, 73,433 people
288 residing in the disaster zone of the Great East Japan Earthquake were analyzed using a cross-sectional
289 design, and the odds ratios for cardiovascular symptoms, such as headache, dizziness, and shortness
290 of breath, were found to be higher for evacuees than for non-evacuees[15]. However, these studies
291 did not consider the combined impact of psychological factors and evacuation status on CVD
292 occurrence.

293 This study identified a higher risk of CVD occurrence among participants having both
294 psychological factors and evacuation status compared with those having neither. Earlier research has
295 demonstrated that disaster-induced psychological burdens and evacuation status are CVD risk
296 factors [12–17]. Earlier studies also identified the loss of social networks as one of the risk factors for
297 CVD [38,39]. In the Health Professionals Follow-up Study, which followed 51,529 American men for
298 10 years, the risk ratio for coronary heart disease was found to be approximately 1.8 times higher for
299 socially isolated men than for men with the highest level of social networks [40]. Hence, the loss of
300 social networks caused by changes in neighbors and a decrease in the frequency of social interaction
301 can lead to future CVD risks. Therefore, the increased CVD risk among people residing in the disaster
302 zone of the Great East Japan Earthquake in Fukushima could have been caused in part by the loss of
303 social networks among participants who were evacuated.

304 Furthermore, the increased risk of CVD due to evacuation status added to psychological factors
305 was observed in men but not in women. To date, some studies after the Great East Japan Earthquake
306 have focused on the association between evacuation status and risk factors for CVD analyzed by
307 gender. In the FHMS, a prospective cohort study from 2011 to 2013 showed an association between
308 evacuation status and hypertension and found the age-adjusted HRs in men to be approximately 1.2
309 times higher than those in women [20]. The same association was observed for overweight, metabolic
310 syndrome, and diabetes, particularly in men [21–23]. These diseases are major risk factors for CVD.
311 Hence, these studies suggest that men might be more susceptible to CVD risk from evacuation status
312 than women following disaster occurrence. Furthermore, symptoms such as hypertension,
313 overweight, metabolic syndrome, or diabetes among male evacuees of the Great East Japan
314 Earthquake might be associated with an increased risk of CVD.

315 This study has the following salient features: (1) identified a large population-based cohort,
316 including 38,710 participants in the affected area immediately after the occurrence of the Great East
317 Japan Earthquake; (2) adopted a longitudinal design (2012–2017); (3) considered the combined impact
318 of psychological factors and evacuation status on both men and women; and (4) adjusted for a range
319 of potential confounders including lifestyle and work-related factors. However, the study also has
320 some limitations. (1) To clarify CVD onset, the study used an annual self-administered survey, whose

overall response rate was not very high (77.1%). However, it considered to have not significantly affect the study's results. In fact, although the study excluded 5,202 men and 6,036 women who never responded to the follow-up surveys, there was no substantial difference in CVD risk factors at baseline between those who responded to follow-up and those who did not. (2) In this study, the definition of evacuation status was based on the living conditions category. Therefore, not all evacuation zones in Fukushima Prefecture designated by the government were included. However, it is considered that the psychological burdens of the actual evacuation life were reflected more accurately by being based on the living conditions category.

5. Conclusions

In summary, this study combined evacuation status with psychological factors, which enabled us to identify a 5% to 25% increased risk of stroke and heart disease in men after the Great East Japan Earthquake in Fukushima. This study provides useful information for healthcare policymakers who plan CVD prevention strategies after major disasters and emphasizes the need for further monitoring of affected areas, particularly with respect to evacuees with psychological burdens. Future research should extend the follow-up period and consider long-term associations in greater detail.

Supplementary Materials: The following are available online at www.mdpi.com/xxx/s1, Table S1: Gender-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to evacuation status.

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