



1 Article

2 **Association between psychological factors and**  
3 **evacuation status and the incidence of cardiovascular**  
4 **diseases after the Great East Japan Earthquake: A**  
5 **prospective study of the Fukushima Health**  
6 **Management Survey**

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

47 men. Evacuation combined with psychological factors increased the risk of stroke and heart disease  
48 risks especially in men after the Great East Japan Earthquake.

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

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## 52 1. Introduction

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

64 There is a complex association between disaster occurrence and CVD risk, and the available  
65 research indicates numerous risk factors, such as lifestyle-related, social, and psychological factors  
66 [10–12]. A number of studies have reported that the disaster was followed by cases of serious post-  
67 traumatic stress disorder (PTSD), which in turn induced CVD in certain individuals affected by the  
68 disaster [13,14]. According to evidence from a few studies, various modifiable risk factors for CVD in  
69 addition to psychological factors have been identified, one of which is evacuation status [15–17]. After  
70 a disaster, the resulting traumatic events and evacuation status affect individuals' lifestyle and  
71 psychological state [18,19]. Earlier studies on the Great East Japan Earthquake have shown that  
72 evacuation status increases the incidence of hypertension, obesity, metabolic syndrome, diabetes,  
73 hyperlipidemia, liver dysfunction, chronic kidney disease, and polycythemia, all of which are CVD  
74 risk factors [20–27]. Although a number of studies have focused on the relationship between  
75 psychological factors and CVD risk or between evacuation status and CVD risk, they have not  
76 clarified how evacuation status affects the association between psychological factors and CVD  
77 occurrence.

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

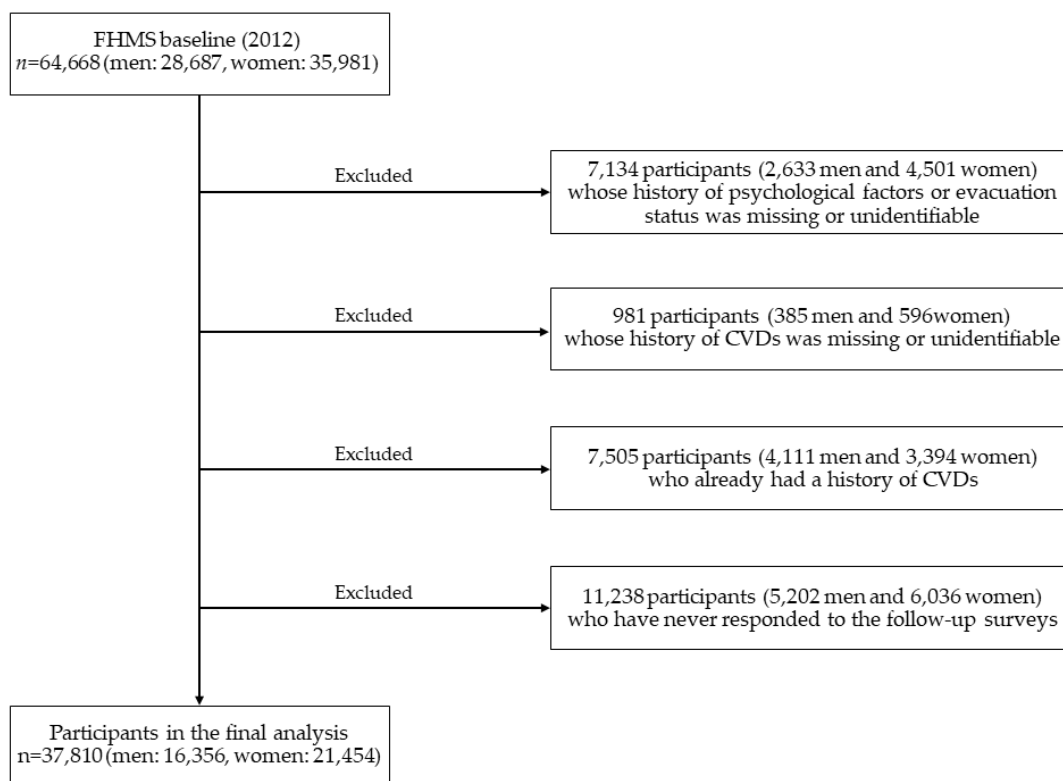
## 85 2. Materials and Methods

### 86 2.1. Participants

87

88 To monitor the health status of Fukushima evacuees after the earthquake and to provide them with  
89 appropriate care, the Fukushima Medical University has been conducting an annual self-administered  
90 survey titled the "Mental Health and Lifestyle Survey [1]." One of its baseline surveys was conducted  
91 in January 2012 to investigate CVD onset, psychological factors, evacuation status, and lifestyle-related  
92 factors. Figure 1 provides a flow chart of the participant selection for the present study. In the survey,  
93 64,668 Japanese individuals (28,687 men, 35,981 women) aged 30–89 years participated from 13  
94 municipalities in the Fukushima Prefecture. Follow-up surveys assessing CVD onset were  
95 subsequently conducted once a year from baseline, and questionnaires were mailed to the participants

96 annually. The present study excluded 2,633 men and 4,501 women whose history of psychological  
 97 factors or evacuation status in 2012 was missing or unidentifiable, as well as 385 men and 596 women  
 98 whose CVDs history in 2012 was missing or unidentifiable. The study also excluded 4,111 men and  
 99 3,394 women who already had a history of CVDs in 2012. The study further excluded 5,202 men and  
 100 6,036 women who never responded to the follow-up surveys. The final analysis therefore included  
 101 37,810 participants (16,356 men, 21,454 women).  
 102  
 103



104  
 105 **Figure 1.** Flow chart of participant selection for the present study of association between  
 106 psychological factors and evacuation status and the incidence of cardiovascular diseases after the  
 107 Great East Japan Earthquake  
 108

109 The study was conducted based on the provisions of the Declaration of Helsinki, and the study  
 110 protocol was approved by the Ethics Review Committees of Okayama University (No. 1803-022) and  
 111 Fukushima Medical University (No. 1316, No. 2148, No. 2020-047). Submission of the self-administered  
 112 questionnaires was considered as consent given by the participants for participating in the study.  
 113

## 114 2.2. Measurements

### 115 2.2.1. Onset and mortality of cardiovascular diseases

116 The study considered both onset and death as cardiovascular incidents. To measure the onset of  
 117 CVDs (stroke or heart disease), the participants were presented once a year with the question, "Have  
 118 you ever been diagnosed by a doctor with the following diseases?" and were asked to circle either "yes"  
 119 or "no" for any applicable diseases [15,28–30]. The mortality data were forwarded to the public health  
 120 departments of the respective areas before being centralized at the Ministry of Health, Labour and  
 121 Welfare, and the underlying causes of death were coded based on the 10th revision of the International  
 122 Classification for Diseases (ICD-10). The primary endpoints for the current analysis were death from  
 123 stroke (ICD-10 codes I60–I69), heart disease (ICD-10 codes I20–I25 and I30–I52), and total CVD (stroke  
 124 and heart disease).  
 125  
 126

### 127 2.2.2. Psychological factors and evacuation status

128 To assess the participants' mental health status, we used the six-item Kessler Psychological  
129 Distress Scale (K6) and the Post-Traumatic Stress Disorder (PTSD) Checklist—Stressor-Specific Version  
130 (PCL-S). In particular, the K6 was employed to measure psychological distress and screen individuals  
131 for nonspecific serious mental illness [31]. The scale included questions on whether the participants had  
132 experienced any of the following six symptoms during the past 30 days: "feeling so sad that nothing  
133 could cheer you up," "feeling nervous," "feeling hopeless," "feeling restless or fidgety," "feeling  
134 everything was an effort," and "feeling worthless." Each question was scored on a five-point Likert-  
135 type scale, with values ranging from 0 to 4. Scores ranged from 0 to 24, and higher scores indicated  
136 lower mental health status. The study validated and employed the Japanese version of K6 [32,33]. We  
137 defined psychological distress as scores  $\geq 13$ .

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

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

### 149 2.2.3. Lifestyle behaviors and social factors

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

166 The study considered job loss, loss of family members, and a high perception of radiation risks to  
167 be social factors. The "job loss" and "loss of family members" categories included the following "yes"  
168 or "no" questions: "Did you become unemployed?" and "Did you lose loved ones in this earthquake?"  
169 A "yes" response to these questions was considered to be confirmation of job loss and loss of family  
170 members. Furthermore, for the "high perception of radiation risks" category, participants answered the  
171 following multiple-choice questions: "How likely do you think acute health problems (e.g., death  
172 within a month) will occur from the current radiation exposure?," "How likely do you think health  
173 problems (e.g., cancer) will occur in the coming years due to the current radiation exposure?," and "To  
174 what extent do you think the current radiation exposure will affect future generations (children,  
175 grandchildren, etc.?)" The participants who answered "3" or "4" (where 1 corresponded to a low  
176 likelihood and 4 to a high likelihood) to any of these questions were considered to have a high  
177 perception of radiation risks.  
178

179

180 *2.3. Statistical Analysis*

181 The age-adjusted mean and prevalence of baseline variables of interest were compared between  
 182 participants according to evacuation status, using analysis of covariance and logistic regression models  
 183 [35]. Furthermore, *p* for differences were calculated between evacuees and non-evacuees. The number  
 184 of cohorts was sufficient for the current analysis, and the risk of stroke was more significant among  
 185 men; hence, the analyses in this study were stratified by gender.

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

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

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

208 **3. Results**

209 During the 3.7-year mean follow-up period, the onset of all CVDs was reported for 2,829 (1,511  
 210 men and 1,318 women), including 626 stroke onset cases (361 men and 265 women); 2,203 heart  
 211 disease onset cases (1,150 men and 1,053 women); and the death of 171 participants from all CVDs  
 212 (86 men and 85 women), including 72 stroke (26 men and 46 women) and 99 heart disease (60 men  
 213 and 39 women) death cases.

214 *3.1. Cardiovascular risk factors of participants at baseline*

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

220 **Table 1.** Age-adjusted mean or prevalence of cardiovascular risk factors for the participants at  
 221 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 $\geq$ once/month (%)	68.6	71.5	<0.001
Physical activity $\geq$ 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
<b>Women</b>			
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 $\geq$ once/month (%)	30.6	71.5	0.38
Physical activity $\geq$ 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

### 222 3.2. Psychological factors or evacuation status on cardiovascular diseases

223 Table 2 depicts the psychological factors or evacuation status-specific age-adjusted and  
 224 multivariable HRs (95% CIs) of CVDs according to gender. For psychological distress, both men  
 225 and women had an increased risk of stroke and heart disease. The respective multivariable HRs  
 226 (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)  
 227 for women. Similarly, for trauma reaction, both men and women showed an increased risk of stroke  
 228 and heart disease. Respective multivariable HRs were 1.78 (1.49–2.14) and 1.43 (1.28–1.59) for men  
 229 and 1.41 (1.17–1.70) and 1.35 (1.22–1.50) for women. For evacuation status, men showed significant  
 230 or borderline significant increased risks for stroke and heart disease. The respective multivariable  
 231 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  
 232 disease based on their evacuation status. There were statistically significant interactions based on  
 233 gender between evacuation status and stroke ( $p = 0.04$ ).

234 **Table 2.** Psychological factors or evacuation status-specific age-adjusted and multivariable HRs  
 235 (95% CIs) of CVDs according to gender.

	Men	Women	$p$ for interaction <sup>a</sup>
Person-years	59,087	79,350	
<b>Psychological distress</b>			
Total CVD (n)	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 <sup>b</sup>	1.34 (1.18–1.52)	1.39 (1.25–1.55)	0.90
<b>Stroke (n)</b>			
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
<b>Heart disease (n)</b>			
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 (n)	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

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

### 240 3.3. Combination of psychological factors and evacuation status on cardiovascular diseases

241 Table 3 depicts the gender-specific age-adjusted and multivariable HRs (95% CIs) of CVDs  
 242 according to the combination of psychological factors and evacuation status. Compared to those  
 243 with neither psychological distress nor evacuation status, those with both had increased risks of  
 244 stroke and heart disease, and the increased magnitude due to evacuation was approximately 0.21  
 245 for stroke and 0.05 for heart disease in men. The respective multivariable HRs (95% CIs) were 1.75  
 246 (1.26–2.44) for stroke and 1.49 (1.22–1.82) for heart disease. Compared to those with neither trauma  
 247 reaction nor evacuation status, those with both had increased risks of stroke and heart disease, and  
 248 the increased magnitude due to evacuation was approximately 0.25 for stroke and about 0.19 for  
 249 heart disease in men. The respective multivariable HRs were 2.01 (1.54–2.61) for stroke and 1.57  
 250 (1.34–1.84) for heart disease. Evacuation status in addition to psychological factors did not lead to  
 251 an additional impact on CVDs in women.

252 **Table 3.** Gender-specific age-adjusted and multivariable HRs (95% CIs) of CVDs according to the  
 253 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 <sup>a</sup>	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)

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

256 Supplementary Table S1 depicts the gender-specific age-adjusted and multivariable HRs (95% CIs)  
 257 of CVDs according to evacuation status. Men with psychological distress showed an increased risk  
 258 of both stroke and heart disease, irrespective of evacuation status. The respective multivariable HRs  
 259 (95% CIs) were 1.49 (1.07–2.08) for evacuees and 1.54 (1.13–2.11) for non-evacuees for stroke, and  
 260 1.39 (1.14–1.69) vs 1.41 (1.18–1.70) for heart disease. Similarly, men with trauma reaction showed an  
 261 increased risk, and respective multivariable HRs of 1.79 (1.36–2.35) vs 1.75 (1.37–2.24) for stroke,  
 262 and 1.52 (1.29–1.79) vs 1.35 (1.16–1.57) for heart disease. Women with psychological distress  
 263 showed a significant or borderline significant increased risk of both stroke and heart disease,  
 264 irrespective of evacuation status. The respective multivariable HRs (95% CIs) were 1.50 (1.07–2.11)



265 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)  
266 for heart disease. Women with a trauma reaction showed an increased risk of stroke for non-  
267 evacuees and heart disease irrespective of evacuation status. The respective multivariable HRs (95%  
268 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  
269 heart disease.

#### 270 4. Discussion

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

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

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

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

316 This study has the following salient features: (1) identified a large population-based cohort,  
317 including 38,710 participants in the affected area immediately after the occurrence of the Great East  
318 Japan Earthquake; (2) adopted a longitudinal design (2012–2017); (3) considered the combined impact  
319 of psychological factors and evacuation status on both men and women; and (4) adjusted for a range  
320 of potential confounders including lifestyle and work-related factors. However, the study also has  
321 some limitations. (1) To clarify CVD onset, the study used an annual self-administered survey, whose  
322 overall response rate was not very high (77.1%). However, it considered to have not significantly  
323 affect the study's results. In fact, although the study excluded 5,202 men and 6,036 women who never  
324 responded to the follow-up surveys, there was no substantial difference in CVD risk factors at  
325 baseline between those who responded to follow-up and those who did not. (2) In this study, the  
326 definition of evacuation status was based on the living conditions category. Therefore, not all  
327 evacuation zones in Fukushima Prefecture designated by the government were included. However,  
328 it is considered that the psychological burdens of the actual evacuation life were reflected more  
329 accurately by being based on the living conditions category.

## 330 5. Conclusions

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

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

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340 F.H., E.E. and T.O.; investigation and data curation, T.O., M.M., S.Y., Y.S., H.Y., A.T., K.T. M.H. and K.K.;  
341 resources, T.O., M.M., S.Y., Y.S., H.Y., A.T., K.T., M.H., T.H., K.O., H.K. and K.K.; writing and visualization, T.S.  
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