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Community social support and onset of dementia in older Japanese individuals: A multilevel analysis using the JAGES cohort data

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Community social support and onset of dementia in older Japanese individuals: A multilevel analysis using the JAGES cohort data

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ABSTRACT

Objective: Recently, there has been an increase in the number of people with dementia. However, no study has examined the association between community-level social support and the onset of incident dementia using multilevel survival analysis.

Design: A prospective cohort study.

Participants and setting: We analyzed data pertaining to 15,313 community-dwelling adults aged 65 years or older (7,381 men, 7,932 women) who had not accessed long-term care insurance and were living in Aichi Prefecture (seven municipalities) in Japan.

Primary and secondary outcome measures: The association between community-level social support and onset of incident dementia was examined using the Japan Gerontological Evaluation Study, a prospective cohort study introduced in Japan in 2003. Incident dementia was assessed using Long-term Care Insurance records spanning 3,436 days from the baseline survey.

Results: During the 10-year follow-up, the onset of incident dementia occurred in 1,776 adults. Among older people, a 1% increase in community-level social support (in the form of receiving emotional support) was associated with an approximately 4% reduction in the risk of developing dementia, regardless of socio-demographic variables and health conditions (HR = 0.96; 95% CI = 0.94-0.99).

Conclusions: Receiving community-level social support in the form of emotional support is associated with a lower risk of developing incident dementia.

Keywords: Cognitive decline, Population health, Social epidemiology

STRENGTHS AND LIMITATIONS OF THIS STUDY

- To date, no study has examined the association between community-level social support and the onset of incident dementia using multilevel survival analysis.
- This is a long-term follow-up study that followed older adults in Japan for about 10 years.
- The sample does not fully reflect the older population in Japan because the study subjects were recruited from a single prefecture.

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INTRODUCTION

Dementia constitutes a pressing health challenge, especially among the older population. The incidence of dementia worldwide is projected to rise to 66 and 131 million by 2030 and 2050, respectively.[1] In Japan alone, it is predicted that there will be 4.62 and 7 million people affected by 2012 and 2025, respectively. These rates suggest that about one in seven Japanese people aged 65 years or above may develop dementia.[2]

Currently, no effective therapeutic intervention for dementia has been determined. As such, identifying adjustable risks and preventive measures is essential for slowing down or preventing the onset of dementia.[3] Previous studies have identified genetic, vascular, and lifestyle-related factors,[4-9] such as advanced age, being female, having a low education level, being in poor health, smoking, and heavy drinking, as being associated with a higher risk of developing dementia. An additional significantly adjustable risk factor is the lack of positive social networks and influences. A previous study suggested that engagement in social activities, and having a rich network of activities within close relationships, confers some protection against dementia among older people.[4]

The definition and the attendant use of the notion of social relationships vary among researchers. The concept may encompass factors such as social participation, social networks, and social support. Social support might also be a significant protective factor for cognitive aging.[10] Social support has been categorized into four types: giving and receiving support at an emotional level, as well as giving and receiving support at an instrumental level.[11] These types of support occur at an individual level and have been associated with improved health. For example, providing emotional and instrumental social support to non-family members leads to fewer depressive symptoms.[12] Providing emotional support to relatives, friends, and neighbors, along with instrumental support to spouses, further lowers the risk of mortality.[13] Receiving emotional support is associated with improved cognitive

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function.[10] However, incident dementia has been reported in instances where social support is extended between co-resident members of the family.[13]

Besides individual-level social factors, social networks, and relationships at the community level have been investigated to understand its association with moderating the risk of functional disability. Such studies investigate "social capital," that has been defined as the "resources that are accessed by individuals as a result of their membership of a network or a group."[14] A previous study found that lower social capital at the community-level is associated with an increased rate of functional disability among older women.[11] And there are reports of research on community social capital and cognitive decline.[15] However, because these studies are cross-sectional studies, they do not take into account temporal pre/posterior relationships. Nevertheless, no studies have examined how community-level social support influences the risk of dementia. Therefore, this study seeks to evaluate the relationship between social support at the community level and the onset of dementia.

METHODS

Sample

Data for this study were accessed via the Japan Gerontological Evaluation Study (JAGES). Set up in Aichi in 2003, this is a prospective cohort study of the Center for Well-being and Society of the Nihon Fukushi University.[16] The research was carried out in seven municipalities encompassing the entire southern region of the Chita peninsula and the Aichi Prefecture. In October 2003, an estimated 276,208 people resided in these locations. Of this population, 18.0% were aged 65 years or above.[14] On average, the data of 6,300 residents in the 44 school districts were analyzed in this study. Care was taken to limit the sample to people aged 65 years and above who were not recipients of long-term public health insurance benefits due to physical or mental challenges. A random selection of 33,152 people aged 65

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years or older was performed. Of this sample, 15,313 individuals answered the baseline survey (response rate = 52.1%).[13] The exclusion criteria were: a) people who had difficulty in performing activities of daily living as a result of disabilities; b) people who did not provide baseline information (n=579); c) people who did not provide social support information (n = 1359); and d) people did not provide a school district code (n = 2,343).

Ultimately, 11,032 subjects (5,627 women and 5,405 men) were included in this analysis.

The ethics committee of Research of Human Subjects at the Nihon Fukushi University evaluated and approved the use of the JAGES protocol (approval number 13-14). Consent to participate in the study was indicated by a written explanation at the beginning of the questionnaire and by the response received on the questionnaire.

Follow-up

The evaluation parameters of the JAGES Project included health status, functional deterioration, and mental impairment amongst older Japanese people who were not institutionalized. In Japan, there is a long-term care insurance system that covers both institutional and community-based caregiving. Individuals aged 65 years or above qualify to receive benefits on the strict basis of physical and cognitive disability. The follow-up began on November 1, 2003. Dementia-associated data from the six municipalities (specifically in terms of the onset) was assessed until March 28, 2013.

Outcome variables

Dementia was graded on a scale that includes categories from I to IV, and M based on the Activities of Daily Living Independence Assessment Criteria for Older Individuals with Dementia. The Degree of Autonomy in the Daily Lives of Older Individuals with Dementia Scale, created by the Ministry of Health, Labor, and Welfare of Japan, evaluates an

individual's ability to carry out daily tasks associated with living on a scale that includes categories from I to IV and M.

This scale was validated based on its high association with the Mini-Mental State Evaluation.[13, 17] A score of I indicates that the patient suffers from some level of cognitive decline but remains able to perform domestic and social tasks nearly independently. A score of II indicates that the patient has certain symptoms or behaviors indicative of cognitive impairment and challenges in communication that may hamper the performance of daily tasks, although some amount of external assistance is needed to facilitate routine function. A score of III indicates that the patient periodically exhibits symptoms indicating communication challenges or symptoms/behaviors, which may interfere with the performance of daily tasks, necessitating external assistance. A score of IV indicates that the patient usually shows communication or behavioral challenges, which hampers performing daily tasks, necessitating frequent care. Finally, a score of M is used when the patient shows significant cognitive impairment, displays difficult behavior, or has a serious physical illness, requiring expert medical intervention.[18, 19]

Explanatory variables

 An aggregate of individual-level background data was acquired for the 44 school-based districts to evaluate community social support. An aggregate of responses for individual-level social support among the school districts was used as an indicator of community social support. Within the Japanese context, school districts (or primary schools) are primary residential units of individuals within rural zones. Generally, school districts comprise geographical settings where older individuals may readily travel via foot or bicycle.[20] Individual-level social support was assessed based on four dimensions of the Two-Way Social Support Scale.[12] The four types of support included: (a) receiving support at the

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emotional level, (b) providing support at the emotional level, (c) receiving support at the instrumental level, and (d) providing support at the instrumental level.

A single item measured each support, "If you or others required additional daily assistance, who would you depend upon to assist or to be assisted by?" Receiving emotional support was conceptualized as the perception of the respondent's complaints or fears by an individual (e.g., "Do you have someone who listens to your concerns and complaints? Circle all that apply. Options included family living together, separated children and relatives, acquaintance/friends/neighbors"). Providing emotional support was conceptualized as the expression of complaints or fears by an individual to the respondents (e.g., "Do you listen to someone's concerns or complaints? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintances/ friends/ neighbors"). Receiving instrumental support was conceived as the rendering of care to the respondent by an individual, if the respondent were ill for many days (Question: "Do you have someone who looks after you when you are sick and confined to a bed for a few days? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors"). Providing instrumental support was defined as nursing of an individual by the respondent if they were ill for many days (e.g., "Do you look after someone when he/she is sick and confined to a bed for a few days? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors").

The percentage of people who responded to each item was considered while determining the level of social support. An aggregation of the responses to the survey items, apropos the four dimensions of social support, was performed for the 44 local districts and further, considered community social support indicators.[12]

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Covariates

Other explanatory variables included: gender (female, male), age (65–69, 70–74, 75–79, 80– 84, and 85 years or older), living conditions (accompanied/unaccompanied), marital status (married, never married, divorced, widowed, other/missing), education (\geq 13 years, 10–12 years, 6–9 years, <6 years, other/missing), present illness (no, yes, missing), depressive symptoms evaluated by the GDS-15 (Geriatric Depression Scale: no depression 0–4 points, mild depression 5–9 points, depression 10–15 points, missing), smoking status (never, former, current, missing), alcohol consumption (no, do not drink every day, drink every day \leq 35 g/day, drink every day >35 g/day, missing), and individual social support.

Statistical analysis

This prospective study employed multilevel survival analysis. The data of 11,032 people living in 44 local districts were used in this study. The multilevel analysis framework relied on the assumption that the health outcome of individuals is partially affected by the district in which they live. The multilevel model evaluated the change in outcome across districts (random effects) and the influences of community-level factors on the outcome, accounting for specific constituent features (fixed effects). Multilevel survival analysis was employed to compute the hazard ratio (HR) and 95% confidence interval (CI) for the onset of dementia at follow-up. The HR of the social support variable was determined as the 1% variation in the proportion of aggregated social support. For the analyses, each of the four social support indicators at the community level was concurrently adjusted. Furthermore, three sensitivity analyses were conducted, excluding (i) one year, (ii) two years, and (iii) three years after baseline. The STATA SE version 13 (Stata Corp., College Station, TX, USA) was used to conduct the analyses.

Patient and public involvement

The patients and public were not involved in the design, conduct, reporting, or dissemination plans of our research.

RESULTS

During the 9.4-year follow-up period (mean=7.9 person-years; standard deviation [SD] =2.5 person-years), dementia onset was observed in 1,776 individuals (16.1%). Supplementary Table S1 (view as supplementary data online) shows the baseline characteristics and frequency of dementia for every 1,000 people/year. The incidence rate of dementia was higher in those who were female, older, living alone, widowed or divorced, those having less than 6 years of education with an existing illness and with a higher score on GDS-15. It was also higher for those who did not consume alcohol, did not get support at the emotional level, did not offer support at the emotional level, and did not receive help at the instrumental level, compared with each counterpart category.

Table 1 shows the mean, range, median, correlation matrix, and SD of the community-level social support indicators in the 44 districts. Spearman's correlation coefficients ranged from -.11 to .44. The average proportion of people receiving community level emotional support was 89.9%, with a range of 82.7% to 93.5%. The proportion of people receiving community level emotional support was moderately correlated with the proportion of people receiving instrumental support (p = .44).

Table 1 Characteristics and Spearman's correlation coefficient matrices for community level
social support indicators ($N = 44$ school districts)

					Spearm	nan's Rank C	Correlation
	%	SD	Min	Max		Coefficier	nt
					1.	2.	3.
1. Community-level receiving	89.9	2.0	82.7	93 5			
emotional support	07.7 2.0		02.7	10.0			
2. Community-level providing	021	2.2	76 1	00 C	11*		
emotional support	83.1	2.2	/0.1	88.0	- .11 [*]		
3. Community-level receiving	04.0	1.(01.2	07 (44*	00*	
instrumental support	94.0	1.6	91.3	97.6	.44*	.08*	
4. Community-level providing	01.0	0.1	05.6	07.0	0.1	4 1 5	
instrumental support	91.9	2.1	83.6	97.9	01	.41*	.20*

SD, standard deviation; Min, minimum; Max, maximum; * p < .05

The results of the multilevel survival analyses (Model 1) for the onset of incident dementia with three sensitivity analyses models (Model 2, 3, and 4) are shown in Supplementary Table S2 (view as supplementary data online). Regarding community-level social support, in Model 1, a significant association was observed between the onset of incident dementia and the proportion of people receiving community level emotional support (HR=0.96; 95% CI=0.94-0.99). On the contrary, significant correlations or relationship between the onset of incident dementia and other community-level social support were absent. In Model 2 of a sensitivity analysis (excluding 1 year after baseline), significant correlations between the onset of incident dementia and receiving community-level emotional support (HR=0.97; 95% CI=0.94-0.99) remained. Model 3 (excluding 2 years after baseline)

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and Model 4 (excluding 3 years after baseline) showed similar results to Models 1 and 2. Regarding individual-level social support, in Model 1, the incidence of dementia was significantly associated with community-level emotional support (HR=0.83; 95% CI=0.73-0.94) as well as instrumental support (HR=0.83; 95% CI=0.73-0.94).

DISCUSSION

To our knowledge, this is the first study to evaluate community-level social support using multilevel survival analysis to investigate the onset of dementia in a large sample of older community-dwelling individuals. There was a prospective association between living in a community with a higher level of social support and a lower occurrence of dementia during the 10-year study period. However, only one of the community-level social support indicators was significantly associated with dementia onset. The outcome of this research has significant implications for public health. Among older people, a 1% rise in receiving community-level emotional support correlated with an approximately 4% decrease in the incidence of dementia, irrespective of socio-demographic factors and health circumstances.

For individual-level social support, providing social support was significantly correlated with a lower risk for dementia. A previous study indicated that providing emotional and instrumental support at the individual level might be a risk factor for the onset of depression.[12] A previous study by Murata and colleagues [21] examined the association between individual-level social support and dementia development in a 10-year cohort. The results showed that receipt of support from friends and neighbors was associated with a lower risk of developing dementia for both men and women. Nonetheless, people who were providing social support might be less likely to develop dementia.

In the present study, among the four kinds of community social support, only community-level emotional support affected the onset of incident dementia, even after

 adjustment for individual-level social support. Two reasons might contribute to this finding. First, a community where people receive high emotional support from each other might be a place where older people are less likely to feel lonely. Indeed, loneliness was found to predict dementia in a previous study.[22] Second, in areas rich in receiving community-level emotional support, there may be older people who maintain a good relationship with their children. A previous study indicated that positive experiences of receiving social support from children predicted the onset of dementia.[23] In the current study, social support from children was evaluated through three questions: "To what extent do they actually comprehend your feelings about things?" "How much can you depend on them if you experience a critical problem?" and "How much can you open up to them if you need to talk about your fears?" A place where many people receive emotional support may be a place where people generally have good interpersonal relationships. Therefore, a community-level indicator of receiving emotional support may only be associated with the onset of dementia.

Community social support may be an element of social capital or community-level social relationships. For this reason, several plausible pathways between receiving community level emotional support and onset of incident dementia were found in the current study. First, community-level social support may influence people's health by shaping health-associated behaviors. This may be done through faster dissemination of health-related information or by increasing the probability of people taking up healthy standards of behavior and moderating behaviors that have negative effects on health. Second, social support may shape health by enhancing the accessibility of local services and facilities. Social involvement of older people may be fostered by accessing services, including transportation, recreational spaces, and community hubs may foster, thus, restricting or arresting the development of dementia. Third, community social support has the potential to foster good cognitive health by minimizing psychological distress. Fourth, places with higher social

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support at the community level generate greater egalitarian political involvement trends. This may lead to the execution of policies that ensure the safety of community members.

It is critical to mention the possible limitations associated with the present study. First, the response rate to the survey (52.1%) could affect the generalizability of the results. Second, there was no information about the type of dementia diagnoses (for instance, Alzheimer's disease, cerebrovascular dementia, or dementia with Lewy bodies). Third, the sample did not fully reflect the older population in Japan because the study subjects were recruited from a single prefecture. Therefore, the findings cannot be generalized to urban areas or places where the population has distinct characteristics. Finally, other communitylevel social relationships, including social capital, were not evaluated. However, we plan to assess a wider range of community-level factors in subsequent studies.

CONCLUSIONS

The results of this study showed that a higher level of social support at the community level is related to a lower incidence of dementia after adjusting for individual-level social support among older individuals. A community level social support indicator (an aggregated value of receiving emotional support) showed a significant association with dementia onset. The present prospective study suggests that receiving emotional support at the community level may result in a lower level of incident dementia among community-dwelling older individuals in Japan.

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Competing Interests

None of the authors have a conflict of interest to declare.

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Author contributions

YM conceived the research idea, participated in the study design, performed statistical analysis, and prepared the manuscript as the primary author. TT developed the research idea, participated in the study design, cooperated in the statistical analysis, and revised the

manuscript. JA and MS assisted in the data analysis and reviewed the manuscript. TT and YS acquired data, collaborated in statistical analysis, and revised the manuscript. As the lead researcher of the JAGES project, KK helped to conceptualize the study. The final manuscript was read and approved by all authors.

Data sharing statement

Data are available upon reasonable request

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			Incidence	Rate (IR) per 1000	Patients w	vith Dementia
			person-ye	ars	(<i>n</i> = 1,776	5, 16.1%)
Individual-Level variables	п	%	IR	95% CI	n	%
Sex						
Male	5,405	49.0	0.049	[0.045, 0.052]	746	13.8
Female	5,627	51.0	0.062	[0.058,0.066]	1,030	18.3
Age (years)						
65-69	4,065	36.9	0.016	[0.014,0.018]	207	5.1
70-74	3,280	29.7	0.044	[0.040,0.048]	428	13.1
75-79	2,228	20.2	0.090	[0.083,0.098]	548	24.6
80-84	1,012	9.2	0.158	[0.143,0.175]	376	37.2
85+	447	4.1	0.289	[0.253,0.330]	217	48.6
Living Alone						
No	9,959	90.3	0.054	[0.051,0.057]	1,558	15.6
Yes	1,073	9.7	0.073	[0.064,0.083]	218	20.3

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Marital Status								
	Married	7,905	71.7	0.044	[0.042,0.047]	1,038	13.1	
	Widowed or divorced	2,745	24.9	0.089	[0.082,0.096]	658	24.0	
	Never married	190	1.7	0.063	[0.033,0.121]	41	21.6	
	Other/Missing	192	1.7	0.071	[0.052,0.097]	39	20.3	
Educat	ion (years)							
	≥13	455	4.1	0.046	[0.039,0.053]	171	37.6	
	10–12	6,002	54.4	0.048	[0.044,0.052]	963	16.0	
	6–9	3,341	30.3	0.055	[0.052,0.059]	470	14.1	
	<6	1,132	10.3	0.156	[0.134,0.181]	149	13.2	
	Other/Missing	102	0.9	0.081	[0.054,0.121]	23	22.6	
Presen	t Illness							
	No	2,906	26.3	0.040	[0.036,0.045]	354	12.2	
	Yes	7,679	69.6	0.062	[0.059,0.065]	1,348	17.6	
	Missing	447	4.1	0.058	[0.046,0.072]	74	16.6	

GDS-15

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0–4	6,737	61.1	0.043	[0.040,0.045]	857	12.7
5–9	2,234	20.3	0.073	[0.067,0.080]	450	20.1
10–15	644	5.8	0.106	[0.091,0.123]	170	26.4
Missing	1,417	12.8	0.076	[0.068,0.085]	299	21.1
Smoking Status						
Never	6,501	58.9	0.059	[0.056,0.063]	1,129	17.4
Former	2,757	25.0	0.047	[0.042,0.052]	367	13.3
Current	1,396	12.7	0.055	[0.048,0.063]	213	15.3
Missing	378	3.4	0.063	[0.050,0.080]	67	17.7
Alcohol Consumption						
None	7,094	64.3	0.063	[0.059,0.066]	1,268	17.9
Not drink every day	1,513	13.7	0.042	[0.036,0.048]	188	12.4
Drink every day ≤35 g/day	1,769	16.0	0.045	[0.039,0.051]	233	13.2
Drink every day >35 g/day	495	4.5	0.033	[0.025,0.044]	49	9.9
Missing	161	1.5	0.085	[0.062,0.117]	38	23.6
Social Support						

Receiving Emotional Support									
No	1,089	9.9	0.070	[0.061,0.080]	208	19.1			
Yes	9,943	90.1	0.054	[0.052,0.057]	1,568	15.8			
Providing Emotional Support									
No	1,836	16.6	0.085	[0.077,0.094]	409	22.3			
Yes	9,196	83.4	0.051	[0.048,0.053]	1,367	14.9			
Receiving Instrumental Support									
No	868	7.9	0.063	[0.053,0.076]	112	18.1			
Yes	10,164	92.1	0.055	[0.053,0.058]	1,664	16.0			
Providing Instrumental Support									
No	619	5.6	0.113	[0.100,0.128]	250	28.8			
Yes	10,413	94.4	0.051	[0.049,0.054]	1,526	15.0			
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	Model 1	Model 2 (1 year)	Model 3 (2 year)	Model 4 (3 year
	<i>n</i> = 11,032	n = 10,780	<i>n</i> = 10,440	<i>n</i> = 10,071
Fixed Effect	HR (95% CI)	HR (95% CI)	HR (95% CI)	HR (95% CI)
Community-Level Variables				
Rate of receiving emotional support*	0.96 (0.94–0.99)	0.97 (0.94–0.99)	0.97 (0.94–0.99)	0.97 (0.94–0.99
Rate of providing emotional support*	0.99 (0.96–1.01)	0.99 (0.96–1.01)	0.98 (0.95–1.01)	0.98 (0.95-1.01
Rate of receiving instrumental support*	1.01 (0.97–1.04)	1.01 (0.97–1.04)	1.01 (0.97–1.05)	1.01 (0.97–1.06
Rate of providing instrumental				
support*	1.00 (0.97–1.03)	1.00 (0.97–1.02)	1.00 (0.97–1.03)	1.00 (0.97–1.04
Individual-Level Variables				
Social Support				
Receiving Emotional Support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.04 (0.88–1.22)	1.03 (0.87–1.22)	1.03 (0.87–1.23)	0.99 (0.83–1.19
Providing Emotional Support				
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No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.83 (0.73-0.94)	0.88 (0.77–0.99)	0.91 (0.80–1.04)	0.91 (0.79–1.05)
Receiving Instrumental Support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.22 (0.99–1.51)	1.18 (0.95–1.47)	1.13 (0.90–1.42)	1.15 (0.91–1.47)
Providing Instrumental Support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.76 (0.66–0.89)	0.82 (0.70-0.96)	0.85 (0.72–1.01)	0.89 (0.74–1.07)
Sex				
Male	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Female	1.01 (0.87–1.18)	1.03 (0.88–1.20)	1.06 (0.90–1.24)	1.08 (0.91–1.28)
Age (years)				
65–69	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
70–74	2.64 (2.23–3.12)	2.67 (2.25–3.16)	2.65 (2.22–3.16)	2.71 (2.26–3.25)
75–79	5.42 (4.60-6.38)	5.59 (4.73–6.61)	5.84 (4.93-6.93)	5.97 (5.00-7.13)
80–84	9.85 (8.24–11.78)	10.08 (8.40–12.10)	10.30 (8.53–12.43)	10.31 (8.46–12.56)

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85+	19.01 (15.38–23.50)	19.32 (15.52–24.06)	19.32 (15.33–24.34)	18.64 (14.52–23.92)
Living Alone				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.94 (0.80–1.11)	0.94 (0.79–1.12)	0.98 (0.82–1.17)	1.03 (0.86–1.24)
Marital status				
Married	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Widowed or Divorced	1.09 (0.96–1.24)	1.09 (0.96–1.24)	1.06 (0.92–1.21)	1.00 (0.86–1.15)
Never Married	1.28 (0.92–1.78)	1.22 (0.87–1.73)	1.24 (0.87–1.76)	1.25 (0.87–1.79)
Other/Missing	1.12 (0.80–1.56)	1.13 (0.80–1.58)	1.09 (0.77–1.56)	1.09 (0.76–1.57)
Education (years)				
≥13	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
10–12	0.89 (0.74–1.08)	0.89 (0.74–1.08)	0.89 (0.73–1.08)	0.92 (0.75–1.13)
6-9	0.96 (0.80–1.15)	0.95 (0.79–1.14)	0.95 (0.79–1.15)	0.96 (0.79–1.16)
<6	1.29 (1.02–1.63)	1.24 (0.97–1.57)	1.22 (0.95–1.57)	1.23 (0.95–1.61)
Other/Missing	0.89 (0.57–1.40)	0.82 (0.51–1.33)	0.80 (0.48–1.33)	0.86 (0.51–1.45)

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Present Illness

No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.15 (1.02–1.30)	1.13 (1.00–1.28)	1.13 (1.00–1.28)	1.10 (0.96–1.26)
Missing	0.96 (0.75–1.24)	0.98 (0.75–1.27)	0.99 (0.75–1.29)	1.01 (0.76–1.33)
GDS-15				
0-4	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
5-9	1.54 (1.37–1.73)	1.54 (1.37–1.74)	1.48 (1.31–1.68)	1.49 (1.30–1.69)
10-15	2.33 (1.96–2.77)	2.18 (1.82–2.62)	2.16 (1.79–2.62)	2.17 (1.78–2.66)
Missing	1.47 (1.28–1.68)	1.43 (1.25–1.64)	1.41 (1.22–1.63)	1.41 (1.22–1.64)
Smoking Status				
Never	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Former	0.99 (0.85–1.16)	1.00 (0.86–1.18)	0.99 (0.84–1.17)	1.02 (0.86–1.21)
Current	1.34 (1.12–1.60)	1.32 (1.10–1.59)	1.39 (1.15–1.68)	1.40 (1.15–1.71)
Missing	0.86 (0.65–1.15)	0.90 (0.67–1.20)	0.85 (0.63–1.15)	0.91 (0.67–1.24)
Alcohol Consumption				
None	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)

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Does not drink every day				
Does not utnik every duy	0.96 (0.82–1.13)	0.96 (0.81_1.13)	0.97 (0.82–1.15)	0.96 (0.80–1.15
Drinks every day ≤35 g/day	0.98 (0.84–1.15)	0.99 (0.84–1.16)	1.02 (0.86–1.20)	1.03 (0.87–1.22)
Drinks every day >35 g/day	0.86 (0.64–1.17)	0.90 (0.66–1.21)	0.92 (0.68–1.26)	0.91 (0.66–1.26
Missing	1.14 (0.78–1.66)	1.14 (0.78–1.67)	1.24 (0.84–1.83)	1.22 (0.82–1.83)
Random effects				
Community-level variance (SE)	0.06 (0.05)	0.04 (0.06)	0.04 (0.07)	0.09 (0.04)

STROBE Statement-checklist of items that should be included in reports of observational studies

	Item No	Recommendation	Page No
Title and abstract	1	(<i>a</i>) Indicate the study's design with a commonly used term in the title or	1
		the abstract	
		(b) Provide in the abstract an informative and balanced summary of what	2
		was done and what was found	
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4-5
Objectives	3	State specific objectives, including any prespecified hypotheses	5
Methods			
Study design	4	Present key elements of study design early in the paper	5
Setting	5	Describe the setting, locations, and relevant dates, including periods of	5-6
	-	recruitment, exposure, follow-up, and data collection	
Participants	6	(a) Cohort study—Give the eligibility criteria, and the sources and	5-6
i un ore ip units	0	methods of selection of participants. Describe methods of follow-up	
		<i>Case-control study</i> —Give the eligibility criteria and the sources and	
		methods of case ascertainment and control selection. Give the rationale	
		for the choice of cases and controls	
		Cross-sectional study—Give the eligibility criteria and the sources and	
		methods of selection of participants	
		(b) Cohort study — For matched studies, give matching criteria and	5.6
		(b) Conort study—1 of matched studies, give matching effectia and	5-0
		Case control study. For matched studies, give matching criteria and the	
		number of controls per case	
Variables	7	Clearly define all outcomes exposures predictors potential confounders	67
variables	/	and effect modifiers. Give diagnostic criteria if applicable	0-7
Data sources/	8*	For each variable of interest, give sources of data and details of methods	6-9
measurement	0	of assessment (measurement). Describe comparability of assessment	0-7
medsurement		methods if there is more than one group	
Bias	9	Describe any efforts to address notential sources of bias	5-6
Study size	10	Explain how the study size was arrived at	5-6
Quantitative variables	10	Explain how the study size was arrived at	6.0
Quantitative variables	11	applicable, describe which groupings were chosen and why	0-9
Statistical mathada	12	(a) Describe all statistical methods, including these used to control for	6.0
Statistical methods	12	(a) Describe an statistical methods, including those used to control for	0-9
		(b) Describe any methods used to examine subgroups and interactions	6.0
		(b) Describe any methods used to examine subgroups and interactions	6-9
		(c) Explain now missing data were addressed	6-9
		(d) Cohort study—If applicable, explain how loss to follow-up was	6
		addressed	
		<i>Case-control stuay</i> —II applicable, explain now matching of cases and	
		controls was addressed	
		<i>Cross-sectional study</i> —It applicable, describe analytical methods taking	
		account of sampling strategy	
		(<u>e</u>) Describe any sensitivity analyses	9

Continued on next page

Results			
Participants	13*	(a) Report numbers of individuals at each stage of study-eg numbers potentially	5
		eligible, examined for eligibility, confirmed eligible, included in the study,	
		completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	
		(c) Consider use of a flow diagram	
Descriptive	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and	1
data		information on exposures and potential confounders	
		(b) Indicate number of participants with missing data for each variable of interest	
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	6
Outcome data	15*	Cohort study—Report numbers of outcome events or summary measures over time	6
		Case-control study-Report numbers in each exposure category, or summary	
		measures of exposure	
		Cross-sectional study—Report numbers of outcome events or summary measures	
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and	1
		their precision (eg, 95% confidence interval). Make clear which confounders were	1
		adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	1
			1
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a	
		meaningful time period	
Other analyses	17	Report other analyses done-eg analyses of subgroups and interactions, and	1
		sensitivity analyses	1
Discussion			
Key results	18	Summarise key results with reference to study objectives	1
			1
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or	1
		imprecision. Discuss both direction and magnitude of any potential bias	
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations,	1
		multiplicity of analyses, results from similar studies, and other relevant evidence	
Generalisability	21	Discuss the generalisability (external validity) of the study results	1
			1
Other informati	on		
Funding	22	Give the source of funding and the role of the funders for the present study and, if	1
0			

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.

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Community social support and onset of dementia in older Japanese individuals: A multilevel analysis using the JAGES cohort data

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Primary Subject Heading :	Epidemiology
Secondary Subject Heading:	Epidemiology
Keywords:	Dementia < NEUROLOGY, EPIDEMIOLOGY, SOCIAL MEDICINE





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1	Community social support and onset of dementia in older Japanese individuals: A
2	multilevel analysis using the JAGES cohort data
3	
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23	
24	Word count: 3401words

 $\mathbf{2}$

1		2
2 3 4	1	ABSTRACT
5 6	2	Objective : Recently, there has been an increase in the number of people with dementia.
/ 8 0	3	However, no study has examined the association between community-level social support
) 10 11	4	and the onset of incident dementia using multilevel survival analysis.
12 13	5	Design: A prospective cohort study.
14 15	6	Participants and setting: We analyzed data pertaining to 15,313 community-dwelling adults
16 17 18	7	aged 65 years or older (7,381 men, 7,932 women) who had not accessed long-term care
19 20	8	insurance and were living in Aichi Prefecture (seven municipalities) in Japan.
21 22	9	Primary and secondary outcome measures: The association between community-level
23 24 25	10	social support and onset of incident dementia was examined using the Japan Gerontological
26 27	11	Evaluation Study, a prospective cohort study introduced in Japan in 2003. Incident dementia
28 29	12	was assessed using Long-term Care Insurance records spanning 3,436 days from the baseline
30 31 32	13	survey.
32 33 34	14	Results: During the 10-year follow-up, the onset of incident dementia occurred in 1,776
35 36	15	adults. Among older people, a 1% increase in community-level social support (in the form of
37 38	16	receiving emotional support) was associated with an approximately 4% reduction in the risk
39 40 41	17	of developing dementia, regardless of socio-demographic variables and health conditions
42 43	18	(HR = 0.96; 95% CI = 0.94-0.99).
44 45	19	Conclusions: Receiving community-level social support in the form of emotional support is
46 47 48	20	associated with a lower risk of developing incident dementia.
49 50	21	
51 52	22	Keywords: Cognitive decline, Population health, Social epidemiology
53 54 55	23	
55 56 57	24	
58 59 60	25	

1 STRENGTHS AND LIMITATIONS OF THIS STUDY

- To date, no study has examined the association between community-level social support and the onset of incident dementia using multilevel survival analysis.
 - This is a long-term follow-up study that followed older adults in Japan for about 10 years.
- The sample does not fully reflect the older population in Japan because the study subjects were recruited from a single prefecture.

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1 INTRODUCTION

Dementia constitutes a pressing health challenge, especially among the older population. The
incidence of dementia worldwide is projected to rise to 66 and 131 million by 2030 and 2050,
respectively.[1] In Japan alone, it is predicted that there will be 4.62 and 7 million people
affected by 2012 and 2025, respectively. These rates suggest that about one in seven Japanese
people aged 65 years or above may develop dementia.[2]

Currently, no effective therapeutic intervention for dementia has been determined. As such, identifying adjustable risks and preventive measures is essential for slowing down or preventing the onset of dementia.[3] Previous studies have identified genetic, vascular, and lifestyle-related factors, [4-9] such as advanced age, being female, having a low education level, being in poor health, smoking, and heavy drinking, as being associated with a higher risk of developing dementia. An additional significantly adjustable risk factor is the lack of positive social networks and influences. A previous study suggested that engagement in social activities, and having a rich network of activities within close relationships, confers some protection against dementia among older people.[4]

The definition and the attendant use of the notion of social relationships vary among researchers. The concept may encompass factors such as social participation, social networks, and social support. Social support can be defined as aid and assistance exchanged through social relationships and interpersonal transactions [10], and it might be a significant protective factor for cognitive aging.[11] A previous systematic review paper indicated that people with social support had 50% lower mortality than those without it.[12] Social support has been categorized into four types—giving and receiving support at an emotional level and at an instrumental level[13]—all of which occur at an individual level and have been associated with improved health. For example, providing emotional and instrumental social support to non-family members leads to fewer depressive symptoms.[13] Providing

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emotional support to the spouse, and instrumental support to relatives, friends, and neighbors, further lowers the risk of mortality.[14] Receiving emotional support is associated with improved cognitive function.[11] In addition, diverse social relationships, including social support from family members, are associated with a lower incidence of dementia.[15] Besides individual-level social factors, social networks and relationships at the community level have been investigated to understand its association with moderating the risk of functional disability. Such studies investigate "social capital," which has been defined as the "resources that are accessed by individuals as a result of their membership of a network or a group."[16] A previous study found that lower social capital at the community-level is associated with an increased rate of functional disability among older women.[17] Moreover, there are reports of research on community social capital and cognitive decline.[18] However, because these studies are cross-sectional ones, longitudinal studies are needed. Nevertheless, no study has examined how community-level social support influences the risk of dementia. Therefore, this study seeks to evaluate the relationship between social support at the community level and the onset of dementia.

17 METHODS

18 Sample

Data for this study were accessed via the Japan Gerontological Evaluation Study (JAGES).
Set up in Aichi in 2003, this was a prospective cohort study of the Center for Well-being and
Society of the Nihon Fukushi University.[19] The research was carried out in seven—three
larger (Handa, Tokoname, and Chita Hokubu) and four smaller (Agui, Mihama, Minamichita,
and Taketoyo)— municipalities that encompass the entire southern region of the Chita
peninsula and the Aichi Prefecture. In October 2003, an estimated 276,208 people resided in
these locations where 18.0% were aged 65 years or above.[17] On average, the data of 6,300

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2 3 4	1	residents in the 44 school districts were analyzed in this study. From the three larger
5 6	2	municipalities, 5,000 survey candidates were randomly selected from the list of persons
7 8 0	3	insured by long-term care insurance who were not certified as requiring long-term care. In the
9 10 11	4	four smaller municipalities, candidates were randomly selected from those not receiving
12 13	5	public long-term care insurance benefits due to a physical or cognitive disability. Of the
14 15	6	33,152 people selected, 15,313 individuals answered the baseline survey (response rate =
16 17 18	7	52.1%).[15] The exclusion criteria were: a) people who had difficulty in performing activities
19 20	8	of daily living as a result of disabilities; b) people who did not provide baseline information
21 22	9	(n=579); c) people who did not provide social support information ($n = 1359$); and d) people
23 24 25	10	who did not provide a school district code ($n = 2,343$). Ultimately, 11,032 subjects (5,627)
26 27	11	women and 5,405 men) were included in this analysis.
28 29	12	The ethics committee of Research of Human Subjects at the Nihon Fukushi
30 31 22	13	University evaluated and approved the use of the JAGES protocol (approval number 13-14).
32 33 34	14	Consent to participate in the study was indicated by a written explanation at the beginning of
35 36	15	the questionnaire and by the response received on the questionnaire.
37 38	16	
39 40 41	17	Follow-up
42 43	18	The evaluation parameters of the JAGES Project included health status, functional
44 45	19	deterioration, and mental impairment amongst older Japanese people who were not
46 47 48	20	institutionalized. In Japan, there is a long-term care insurance system that covers both
49 50	21	institutional and community-based caregiving. Individuals aged 65 years or above qualify to
51 52	22	receive benefits on the strict basis of physical and cognitive disability. The follow-up began
53 54	23	on November 1, 2003. Dementia-associated data from the six municipalities (specifically in
55 56 57	24	terms of the onset) was assessed until March 28, 2013.
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1	Outcome variables
2	Dementia was graded on a scale that includes categories from I to IV, and M based on the
3	Activities of Daily Living Independence Assessment Criteria for Older Individuals with
4	Dementia. The Degree of Autonomy in the Daily Lives of Older Individuals with Dementia
5	Scale, created by the Ministry of Health, Labor, and Welfare of Japan, evaluates an
6	individual's ability to carry out daily tasks associated with living on a scale that includes
7	categories from I to IV and M (Supplementary Table S1, view as supplementary data online).
8	This scale was validated based on its high association with the Mini-Mental State
9	Evaluation.[20] It has been reported that dementia symptoms indices are strongly correlated
10	with Mini Mental State Examination. (Spearman's rank correlation $\rho = -0.73$, P < 0.001).
11	Scores I, II, III, and IV on the dementia scale are equivalent to 22, 16, 13, and 6 points on the
12	Mini-Mental State Examination, respectively.[20] A score of I indicates that the patient
13	suffers from some level of cognitive decline but remains able to perform domestic and social
14	tasks nearly independently. A score of II indicates that the patient has certain symptoms or
15	behaviors indicative of cognitive impairment and challenges in communication that may
16	hamper the performance of daily tasks, although some amount of external assistance is
17	needed to facilitate routine function. A score of III indicates that the patient periodically
18	exhibits symptoms indicating communication challenges or symptoms/behaviors, which may
19	interfere with the performance of daily tasks, necessitating external assistance. A score of IV
20	indicates that the patient usually shows communication or behavioral challenges, which
21	hampers performing daily tasks, necessitating frequent care. Finally, a score of M

(M=Medical, requires specialized medical care) is used when the patient shows significant
 cognitive impairment, displays difficult behavior, or has a serious physical illness, requiring
 expert medical intervention. Symptoms and behaviors seen in the M rank include delirium,

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paranoia, agitation, self-injury and harm, and other psychiatric symptoms, as well as ongoing
 problem behaviors caused by psychiatric symptoms.

Explanatory variables

An aggregate of individual-level background data was acquired for the 44 school-based districts to evaluate community social support. An aggregate of responses for individual-level social support among the school districts was used as an indicator of community social support. Within the Japanese context, school districts (or primary schools) are primary residential units of individuals within rural zones. Generally, school districts comprise geographical settings where older individuals may readily travel via foot or bicycle.[21] Individual-level social support was assessed based on four dimensions of the Two-Way Social Support Scale.[13] The four types of support included: (a) receiving support at the emotional level, (b) providing support at the emotional level, (c) receiving support at the instrumental level, and (d) providing support at the instrumental level (Supplementary Table S2, view as supplementary data online).

A single item measured each support, "If you or others required additional daily assistance, who would you depend upon to assist or to be assisted by?" Receiving emotional support was conceptualized as the perception of the respondent's complaints or fears by an individual (e.g., "Do you have someone who listens to your concerns and complaints? Circle all that apply. Options included family living together, separated children and relatives, acquaintance/friends/neighbors"). Providing emotional support was conceptualized as the expression of complaints or fears by an individual to the respondents (e.g., "Do you listen to someone's concerns or complaints? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintances/friends/ neighbors"). Receiving instrumental support was conceived as the rendering of care to the

respondent by an individual, if the respondent were ill for many days (e.g., "Do you have someone who looks after you when you are sick and confined to bed for a few days? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors"). Providing instrumental support was defined as nursing of an individual by the respondent if they were ill for many days (e.g., "Do you look after someone when he/she is sick and confined to bed for a few days? Circle the numbers of all the answers that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors").

The percentage of people who responded to each item was considered while determining the level of social support. An aggregation of the responses to the survey items, apropos the four dimensions of social support, was performed for the 44 local districts and further, considered community social support indicators.

Covariates

Other explanatory variables included: gender (female, male), age (65–69, 70–74, 75–79, 80– 84, and 85 years or older), living conditions (accompanied/unaccompanied), marital status (married, never married, divorced, widowed, other/missing), education (≥ 13 years, 10–12) years, 6–9 years, <6 years, other/missing), present illness (no, yes, missing), depressive symptoms evaluated by the GDS-15 (Geriatric Depression Scale: no depression 0-4 points, mild depression 5–9 points, depression 10–15 points, missing), smoking status (never, former, current, missing), alcohol consumption (no, do not drink every day, drink every day \leq 35 g/day, drink every day >35 g/day, missing), and individual social support.

Statistical analysis

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1	This prospective study employed multilevel survival analysis. The data of 11,032 people
2	living in 44 local districts were used in this study. The multilevel analysis framework relied
3	on the assumption that the health outcome of individuals is partially affected by the district in
4	which they live. The multilevel model evaluated the change in outcome across districts
5	(random effects) and the influences of community-level factors on the outcome, accounting
6	for specific constituent features (fixed effects). Multilevel survival analysis was employed to
7	compute the hazard ratio (HR) and 95% confidence interval (CI) for the onset of dementia at
8	follow-up. The HR of the social support variable was determined as the 1% variation in the
9	proportion of aggregated social support. For the analyses, all four social support indicators at
10	the community level and social-demographic factors were concurrently adjusted.
11	Furthermore, three sensitivity analyses were conducted, excluding (i) one year, (ii) two years,
12	and (iii) three years after baseline. The STATA SE version 13 (Stata Corp., College Station,
13	TX, USA) was used for the analysis, and the "stmixed" command was used (the "mestreg"
14	command has become a standard feature in STATA 14).

Patient and public involvement 16

The patients and public were not involved in the design, conduct, reporting, or dissemination 17 plans of our research. 18

20 RESULTS

During the 9.4-year follow-up period (87,232 person-years), dementia onset was observed in 21 1,776 individuals (16.1%). Supplementary Table S3 (view as supplementary data online) 22 23 shows the baseline characteristics and incidence rate of dementia per 1,000 person-years. The incidence rate of dementia was higher in those who were female, older, living alone, 24 25 widowed or divorced, those having less than 6 years of education with an existing illness and

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3 4	1	with a higher score on GDS-15. It was also higher for those who did not consume alcohol,							
5 6	2	did not get support at the emo	tional level	, did not	t offer su	pport at	the emotio	nal level,	and did
7 8	3	not receive help at the instrumental level, compared with each counterpart category.							
9 10 11	4	Table 1 shows the mea	ın, range, m	nedian, c	correlatio	on matrix	, and SD o	f the	
12 13	5	community-level social suppo	rt indicator	s in the	44 distri	cts. Spea	rman's cor	relation	
14 15	6	coefficients ranged from11	to .44. The	average	proport	ion of pe	ople receiv	ing comm	nunity
16 17	7	level emotional support was 8	9.9%, with	a range	of 82.7%	% to 93.5	%. The pro	oportion o	f
18 19 20	8	people receiving community l	evel emotic	onal sup	port was	moderat	ely correla	ted with t	he
21 22 23	9	proportion of people receiving	; instrumen	tal supp	ort (p =	.44).			
24 25									
26 27	10								
28 29	11	Table 1 Characteristics and S	pearman's	correlati	on coeff	icient ma	atrices for o	communit	y level
30 31	12	social support indicators ($N = 44$ school districts)							
32 33 34			Moon	SD.	Min	Mor	Spearman's C	s Rank Corr oefficient	elation
35 36			Ivican	3D	IVIIII	Iviax _	1	2	3
37 38		1. Community level receiving emotional support	89.9	2.0	82.7	93.5			
39 40		2. Community level providing emotional support	83.1	2.2	76.1	88.6	11*		
41 42		3. Community level receiving	94.0	1.6	01.3	07.6	44*	08*	
4⊰		instrumental support	91.0	1.6	91.5	97.0		.00	
44 45		A. Community level providing instrumental support	91.9	2.1	85.6	97.0	01	.41*	.26*
44 45 46	13	4. Community level providing instrumental support SD, standard deviation; M	91.9	2.1 m; Max	85.6	97.9 um; * <i>p</i> <	01	.41*	.26*
44 45 46 47 48 49	13 14	4. Community level providing instrumental support SD, standard deviation; M The results of the mult	91.9 in, minimu ilevel survi	2.1 m; Max val anal	85.6 , maxim	97.9 97.9 um; * <i>p</i> < odel 1) fe	01	.41* t of incide	.26*
44 45 46 47 48 49 50 51	13 14 15	4. Community level providing instrumental support SD, standard deviation; M The results of the mult dementia with three sensitivity	91.9 in, minimu ilevel survi y analyses r	2.1 m; Max val anal nodels (85.6 , maxim lyses (M Model 2	97.0 97.9 um; * <i>p</i> < odel 1) fe	01 <.05 or the onse 4) are show	.41* t of incide n in Table	.26* ent e 2 and
44 45 46 47 48 49 50 51 52 53	13 14 15 16	4. Community level providing instrumental support SD, standard deviation; M The results of the mult dementia with three sensitivity Supplementary Table S4	91.9 in, minimu ilevel survi y analyses r	2.1 m; Max val anal nodels (85.6 , maxim lyses (M Model 2	97.9 97.9 um; * <i>p</i> < odel 1) fa	01 <.05 or the onse 4) are show	.41* t of incide /n in Table	.26* ent e 2 and
44 45 46 47 48 49 50 51 52 53 54 55 55	13 14 15 16 17	A. Community level providing instrumental support SD, standard deviation; M The results of the mult dementia with three sensitivity Supplementary Table S4	91.9 in, minimu ilevel survi y analyses r	2.1 m; Max val anal nodels (85.6 , maxim lyses (M Model 2	97.9 97.9 um; * <i>p</i> < odel 1) f	01 <.05 or the onse 4) are show	.41* t of incide	.26* ent e 2 and
 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 	13 14 15 16 17 18	A. Community level providing instrumental support SD, standard deviation; M The results of the multi- dementia with three sensitivity Supplementary Table S4 Table 2 Results of multilevel	91.9 in, minimu ilevel survi y analyses r survival an	2.1 m; Max val anal nodels (alyses fo	85.6 , maxim lyses (M Model 2	97.9 97.9 um; * <i>p</i> < odel 1) fa	01 <.05 or the onse 4) are show nt dementi	.41* t of incide /n in Table	.26* ent e 2 and
 44 45 46 47 48 49 50 51 52 53 54 55 56 57 58 59 60 	13 14 15 16 17 18	A. Community level providing instrumental support SD, standard deviation; M The results of the mult dementia with three sensitivity Supplementary Table S4 Table 2 Results of multilevel	91.9 in, minimu ilevel survi y analyses r survival an	1.6 2.1 m; Max val anal nodels (alyses fo	85.6 , maxim lyses (M Model 2 or onset	97.9 97.9 um; * <i>p</i> < odel 1) for c, 3, and 6 of incide	01 <.05 or the onse 4) are show nt dementi Model 2	.41* t of incide /n in Tabl a 2 (1 year)	.26*

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n = 11,032	n=10,780
HR (95%CI)	HR (95%CI)
0.96 (0.94-0.99)	0.97 (0.94-0.99)
0.99 (0.96-1.01)	0.99 (0.96-1.01)
1.01 (0.97-1.04)	1.01 (0.97-1.04)
1.00 (0.97-1.03)	1.00 (0.97-1.02)
0.06 (0.05)	0.04 (0.06)
-	n = 11,032 HR (95%CI) 0.96 (0.94-0.99) 0.99 (0.96-1.01) 1.01 (0.97-1.04) 1.00 (0.97-1.03) 0.06 (0.05)

*HR for one-point increment of community social support (range: 0-100)

HR adjusted for sex, age, living alone, marital status, education, present illness, GDS, smoking status, alcohol consumption, receiving emotional support, providing emotional support, receiving instrumental support, and providing instrumental support. (The full version, including individual-level results, is shown in Table S4).

2	Regarding community-level social support, in Model 1, a significant association was
3	observed between the onset of incident dementia and the proportion of people receiving
4	community level emotional support (HR=0.96; 95% CI=0.94-0.99). On the contrary,
5	significant correlations or relationship between the onset of incident dementia and other
6	community-level social support were absent. In Model 2 of a sensitivity analysis (excluding 1
7	year after baseline), significant correlations between the onset of incident dementia and
8	receiving community-level emotional support (HR=0.97; 95% CI=0.94-0.99) remained.
9	Model 3 (excluding 2 years after baseline) and Model 4 (excluding 3 years after baseline)
10	showed similar results to Models 1 and 2 (Supplementary Table S4). Regarding individual-
11	level social support, in Model 1, the incidence of dementia was significantly associated with
12	receiving individual-level emotional support (HR=0.83; 95% CI=0.73-0.94) as well as
13	providing individual-level instrumental support (HR=0.76; 95% CI=0.66-0.89).
14	
15	DISCUSSION
16	To our knowledge, this is the first study to evaluate community-level social support using
17	multilevel survival analysis to investigate the onset of dementia in a large sample of older

community-dwelling individuals. There was a prospective association between living in a community with a higher level of social support and a lower occurrence of dementia during the 10-year study period. However, only one of the community-level social support indicators was significantly associated with dementia onset. The outcome of this research may have significant implications for public health, i.e., by suggesting potential practical implications useful for policymakers, family members, and medical staff. Because previous intervention research indicated that promoting community through salon activity increased social support in the community [22], providing such activities may be a practical solution to prevent the onset of dementia. Among older people, a 1% rise in receiving community-level emotional support correlated with an approximately 4% decrease in the incidence of dementia, irrespective of socio-demographic factors and health circumstances.

For individual-level social support, providing social support was significantly correlated with a lower risk for dementia. A previous study indicated that providing emotional and instrumental support at the individual level might be a risk factor for the onset of depression.[13] A previous study by Murata and colleagues [23] examined the association between individual-level social support and dementia development in a 10-year cohort. The results showed that receipt of support from friends and neighbors was associated with a lower risk of developing dementia for both men and women. Nonetheless, people who were providing social support might be less likely to develop dementia.

In the present study, among the four kinds of community social support, only
community-level emotional support affected the onset of incident dementia, even after
adjustment for individual-level social support. Two reasons might contribute to this finding.
First, a community where people receive high emotional support from each other might be a
place where older people are less likely to feel lonely. Indeed, loneliness was found to predict
dementia in a previous study.[24] Second, because depression was a risk factor for

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1	developing dementia [25], abundant community-level emotional support may mitigate the
2	risk of depression, thereby decreasing the incidence of dementia. [26] In the current study,
3	social support from children was evaluated through three questions: "To what extent do they
4	actually comprehend your feelings about things?" "How much can you depend on them if
5	you experience a critical problem?" and "How much can you open up to them if you need to
6	talk about your fears?" A place where many people receive emotional support may be a place
7	where people generally have good interpersonal relationships. Therefore, a community-level
8	indicator of receiving emotional support may be associated with the onset of dementia.
9	Community social support may be an element of social capital or community-level
10	social relationships. For this reason, several plausible pathways between receiving
11	community level emotional support and onset of incident dementia were found in the current
12	study. First, community-level social support may influence people's health by shaping health-
13	associated behaviors. This may be done through faster dissemination of health-related
14	information or by increasing the probability of people taking up healthy standards of behavior
15	and moderating behaviors that have negative effects on health. Second, social support may
16	shape health by enhancing the accessibility of local services and facilities. Social
17	involvement of older people may be fostered by accessing services, including transportation,
18	recreational spaces, and community hubs may foster, thus, restricting or arresting the
19	development of dementia. Third, community social support has the potential to foster good
20	cognitive health by minimizing psychological distress. Fourth, places with higher social
21	support at the community level generate greater egalitarian political involvement trends. This
22	may lead to the execution of policies that ensure the safety of community members. In
23	addition, according to a systematic review of social capital including studies mainly
24	conducted in western countries, most of the intervention studies in the last two decades have
25	focused on individual-level changes, with a dearth of studies examining community-level

changes.[27] Furthermore, there are few longitudinal studies, even observational studies, that have produced dementia outcomes. Therefore, the results of this study can contribute to the social capital research agenda for developing intervention research at the community level. It is critical to mention the possible limitations associated with the present study. First, the response rate to the survey (52.1%) could affect the generalizability of the results. However, this response rate was higher than the census conducted by the government (41.8% response rate in the postal survey 2020).[28] Second, the dementia outcome is a nationally standardized scale used by public long-term care insurance, but it is not a clinical diagnosis. Third, there was no information about the type of dementia diagnoses (for instance, Alzheimer's disease, cerebrovascular dementia, or dementia with Lewy bodies). Fourth, the sample did not fully reflect the older population in Japan because the study subjects were recruited from a single prefecture. Therefore, the findings cannot be generalized to urban areas or places where the population has distinct characteristics. Finally, other community-level social relationships, including social capital, were not evaluated. However, we plan to assess a wider range of community-level factors in subsequent studies.

17 CONCLUSIONS

The results of this study showed that a higher level of social support at the community level is related to a lower incidence of dementia after adjusting for individual-level social support among older individuals. A community level social support indicator (an aggregated value of receiving emotional support) showed a significant association with dementia onset. The present prospective study suggests that receiving emotional support at the community level may result in a lower level of incident dementia among community-dwelling older individuals in Japan.

2		
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/ 8 0	3	
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2 3	1	Author contributions
4	T	
5 6 7	2	YM conceived the research idea, participated in the study design, performed statistical
7 8 9	3	analysis, and prepared the manuscript as the primary author. Tabuchi T developed the
10 11	4	research idea, participated in the study design, cooperated in the statistical analysis, and
12 13	5	revised the manuscript. JA and MS assisted in the data analysis and reviewed the manuscript.
14 15 16	6	Tsuji T and YS acquired data, collaborated in statistical analysis, and revised the manuscript.
10 17 18	7	As the lead researcher of the JAGES project, KK helped to conceptualize the study. The final
19 20	8	manuscript was read and approved by all authors.
21 22	9	
23 24 25	10	Data sharing statement
26 27	11	Data are available upon reasonable request.
28 29	12	
30 31 22	13	Ethics Statement
32 33 34	14	This study was approved by the Nihon Fukushi University Ethics Committee (approval
35 36	15	number 13-14).
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Ra	nk	Criteria	Examples of observable symptoms or behaviors
0		Independent	
I		Suffers from certain dementia symptoms, but daily living is almost fully independent in the domestic and social	
II		spheres. Manifests some symptoms/behaviors and communication difficulties that may hinder daily activities, but can be independent if someone takes care of them. The above mentioned	Fraguently gets lost on the street, or makes noticeable
	ll a	conditions in II are observed while outside the domestic sphere.	mistakes in matters that the person was previously able to handle, such as shopping, personal administrative tasks, or financial management.
	ll b	The abovementioned conditions in II are also observed in the domestic sphere.	Is unable to manage taking medication or stay alone at home due to an inability to answer the phone or the door.
III		communication difficulties or symptoms/behaviors that hinder daily activities, thus requiring care.	
	IIIa	Manifests above- mentioned conditions described in III predominantly during the day.	Has difficulty or takes time to change clothes, take meals, defecate, or urinate; puts objects into the mouth, picks up and collects objects, is incontinent, makes loud and incoherent screams, carelessly handles fire, or engages in unhygienic acts or inappropriate sexual acts, etc.
	Шb	Manifests above- mentioned conditions described in III predominantly at night.	Same as rank IIIa.
IV		difficulties communicating or symptoms/behaviors that hinder daily activities and constantly requires care.	Same as rank III.
M*		Manifests significant mental symptoms, problematic behaviors, or severe physical illnesses and requires specialized medical care.	Shows continued mental symptoms, such as delirium, delusions, and agitation, and manifests associated problematic behaviors, such as self-mutilation or harm to others.

Table S1 Criteria of levels of dementia symptomatology in the Japanese LTCI system

*M=Medical

Table 52 Four categories of social support (including questionnane nems)					
	Receive	Provide			
	(a) receiving support at the emotional level	(b) providing support at the emotional level			
Emotional social support	(Question: "Do you have someone who listens to your concerns and complaints? Circle all that apply. Options included family living together, separated children and relatives, acquaintance/friends/neighbors")	(Question: "Do you listen to someone's concerns or complaints? Circle all that apply. Options include family living together, separated children and relatives, acquaintances/friends/neighbors")			
Instrumental social support	(c) receiving support at the instrumental level (Question: "Do you have someone who looks after you when you are sick and confined to bed for a few days? Circle all that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors")	(d) providing support at the instrumental level (Question: "Do you look after someone when he/she is sick and confined to bed for a few days? Circle all that apply. Options include family living together, separated children and relatives, acquaintance/friends/neighbors")			

 Table S2 Four categories of social support (including questionnaire items)

Table S3 Descriptive characteristics of the respondents (n = 11,032)

			Patients Demen [n = 1,776 (with itia 16.1%)]		Incide per 10	nce rate(IR))00 person- years
Individual level variables	n	%	n	%	Person- year	IR	95% CI
Sex							
Male	5,405	49.0	746	13.8	41,871	17.8	[16.6, 19.1]
Female	5,627	51.0	1,030	18.3	45,362	22.7	[21.4, 24.1]
Age (years)							
65-69	4,065	36.9	207	5.1	35,143	5.9	[5.1, 6.7]
70-74	3,280	29.7	428	13.1	26,842	15.9	[14.5,17.5]
75-79	2,228	20.2	548	24.6	16,689	32.8	[30.2,35.7]
80-84	1,012	9.2	376	37.2	6,502	57.8	[52.3,64.0]
85+	447	4.1	217	48.6	2,057	105.5	[92.4,120.5]
Living alone							
No	9,959	90.3	1,558	15.6	79,038	19.7	[18.8,20.7]
Yes	1,073	9.7	218	20.3	8,194	26.6	[23.3,30.4]
Marital status							
Married	7,905	71.7	1,038	13.1	63,992	16.2	[15.3,17.2]
Widowed or divorced	2,745	24.9	658	24.0	20,274	32.5	[30.1,35.0]
Never married	190	1.7	41	21.6	1,453	28.2	[20.8,38.3]
Other/Missing	192	1.7	39	20.3	1,514	25.8	[18.8,35.3]
Education (years)							
≧13	455	4.1	171	37.6	3,004	56.9	[49.0,66.1]
10-12	6,002	54.4	963	16.0	47,576	20.2	[19.0,21.6]
6-9	3,341	30.3	470	14.1	26,908	17.5	[16.0,19.1]
<6	1,132	10.3	149	13.2	8,965	16.6	[14.2,19.5]
Other/Missing	102	0.9	23	22.6	780	29.5	[19.6,44.4]
Present illness							
No	2,906	26.3	354	12.2	24,074	14.7	[13.3,16.3]
Yes	7,679	69.6	1,348	17.6	59,636	22.6	[21.4,23.8]
Missing	447	4.1	74	16.6	3,523	21.0	[16.7,26.4]
GDS-15							

0-4	6,737	61.1	857	12.7	55,179	15.5	[14.5,16.6]
5-9	2,234	20.3	450	20.1	16,840	26.7	[24.4,29.3]
10-15	644	5.8	170	26.4	4,393	38.7	[33.3,45.0]
Missing	1,417	12.8	299	21.1	10,820	27.6	[24.7,31.0]
Smoking status							
Never	6,501	58.9	1,129	17.4	52,412	21.5	[20.3,22.8]
Former	2,757	25.0	367	13.3	21,375	17.2	[15.5,19.0]
Current	1,396	12.7	213	15.3	10,543	20.2	[17.7,23.1]
Missing	378	3.4	67	17.7	2,902	23.1	[18.2,29.3]
Alcohol consumption							
Non	7,094	64.3	1,268	17.9	55,456	22.9	[21.6,24.2]
Does not drink every_day	1,513	13.7	188	12.4	12,275	15.3	[13.3,17.7]
Drinks every_day ≦ 35 g/day	1,769	16.0	233	13.2	14,247	16.4	[14.4,18.6]
Drinks every_day > 35 g/day	495	4.5	49	9.9	4,032	12.2	[9.2,16.1]
Missing	161	1.5	38	23.6	1,222	31.1	[22.6,42.7]
Social supports							
Receiving emotional support							
No	1,089	9.9	208	19.1	8,161	25.5	[22.2,29.2]
Yes	9,943	90.1	1,568	15.8	79,071	19.8	[18.9,20.8]
Providing emotional support							
No	1,836	16.6	409	22.3	13,135	31.1	[28.3,34.3]
Yes	9,196	83.4	1,367	14.9	74,097	18.4	[17.5,19.5]
Receiving instrumental support							
No	868	7.9	112	18.1	4,849	23.1	[19.2,27.8]
Yes	10,164	92.1	1,664	16.0	82,384	20.2	[19.3,21.2]
Providing instrumental support							
No	619	5.6	250	28.8	6,044	41.4	[36.5,46.8]
Yes	10,413	94.4	1,526	15.0	81,189	18.8	[17.9,19.8]

	Model 1	Model 2 (1 year)	Model 3 (2 year)	Model 4 (3 year)
	n = 11,032	n=10,780	n = 10,440	n = 10,071
Fixed effect	HR (95% CI)	HR (95% CI)	HR (95% CI)	HR (95% CI)
Community level variables				
Rate of receiving emotional support*	0.96 (0.94- 0.99)	0.97 (0.94- 0.99)	0.97 (0.94- 0.99)	0.97 (0.94- 0.988)
Rate of providing emotional support*	0.99 (0.96- 1.01)	0.99 (0.96- 1.01)	0.98 (0.95- 1.01)	0.98 (0.95-1.01)
Rate of receiving instrumental support*	1.01 (0.97- 1.04)	1.01 (0.97- 1.04)	1.01 (0.97- 1.05)	1.01 (0.97-1.06)
Rate of providing instrumental support*	1.00 (0.97- 1.03)	1.00 (0.97- 1.02)	1.00 (0.97- 1.03)	1.00 (0.97-1.04)
Individual level variables				
Social supports Receiving emotional support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.04 (0.88- 1.22)	1.03 (0.87- 1.22)	1.03 (0.87- 1.23)	0.99 (0.83-1.19)
Providing emotional support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.83 (0.73- 0.94)	0.88 (0.77- 0.997)	0.91 (0.80- 1.04)	0.91 (0.79-1.05)
Receiving instrumental support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.22 (0.99- 1.51)	1.18 (0.95- 1.47)	1.13 (0.90- 1.42)	1.15 (0.91-1.47)
Providing instrumental support				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.76 (0.66- 0.89)	0.82 (0.70- 0.96)	0.85 (0.72- 1.01)	0.89 (0.74-1.07)
Sex				
Male	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Female	1.01 (0.87- 1.18)	1.03 (0.88- 1.20)	1.06 (0.90- 1.24)	1.08 (0.91-1.28)
Age (years)				
65-69	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
70-74	2.64 (2.23- 3.12)	2.67 (2.25- 3.16)	2.65 (2.22- 3.16)	2.71 (2.26-3.25)
75-79	5.42 (4.60- 6.38)	5.59 (4.73- 6.61)	5.84 (4.93- 6.93)	5.97 (5.00-7.13)
80-84	9.85 (8.24- 11.78)	10.08 (8.40- 12.10)	10.30 (8.53- 12.43)	10.31 (8.46- 12.56)

Table S4 Results of multilevel survival analyses for onset of incident dementia (Full version of Table 3, including individual-level results)

85+	19.01 (15.38- 23.50)	19.32 (15.52- 24.06)	19.32 (15.33- 24.34)	18.64 (14.52- 23.92)
Living alone	20.00)	21.00)	21.31)	23.72)
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	0.94 (0.80- 1.11)	0.94 (0.79- 1.12)	0.98 (0.82- 1.17)	1.03 (0.86-1.24)
Marital status				
Married	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Widowed or divorced	1.09 (0.96- 1.24)	1.09 (0.96- 1.24)	1.06 (0.92- 1.21)	1.00 (0.86-1.15)
Never married	1.28 (0.92- 1.78)	1.22 (0.87- 1.73)	1.24 (0.87- 1.76)	1.25 (0.87-1.79)
Other/Missing	1.12 (0.80- 1.56)	1.13 (0.80- 1.58)	1.09 (0.77- 1.56)	1.09 (0.76-1.57)
Education (years)				
≧13	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
10-12	0.89 (0.74- 1.08)	0.89 (0.74- 1.08)	0.89 (0.73- 1.08)	0.92 (0.75-1.13)
6-9	0.96 (0.80- 1.15)	0.95 (0.79- 1.14)	0.95 (0.79- 1.15)	0.96 (0.79-1.16)
<6	1.29 (1.02- 1.63)	1.24 (0.97- 1.57)	1.22 (0.95- 1.57)	1.23 (0.95-1.61)
Other/Missing	0.89 (0.57- 1.40)	0.82 (0.51- 1.33)	0.80 (0.48- 1.33)	0.86 (0.51-1.45)
Present illness				
No	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Yes	1.15 (1.02- 1.30)	1.13 (1.00- 1.28)	1.13 (1.00- 1.28)	1.10 (0.96-1.26)
Missing	0.96 (0.75- 1.24)	0.98 (0.75- 1.27)	0.99 (0.75- 1.29)	1.01 (0.76-1.33)
GDS-15				
0-4	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
5-9	1.54 (1.37- 1.73)	1.54 (1.37- 1.74)	1.48 (1.31- 1.68)	1.49 (1.30-1.69)
10-15	2.33 (1.96- 2.77)	2.18 (1.82- 2.62)	2.16 (1.79- 2.62)	2.17 (1.78-2.66)
Missing	1.47 (1.28- 1.68)	1.43 (1.25- 1.64)	1.41 (1.22- 1.63)	1.41 (1.22-1.64)
Smoking status				
Never	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Former	0.99 (0.85- 1.16)	1.00 (0.86- 1.18)	0.99 (0.84- 1.17)	1.02 (0.86-1.21)
Current	1.34 (1.12- 1.60)	1.32 (1.10- 1.59)	1.39 (1.15- 1.68)	1.40 (1.15-1.71)
Missing	0.86 (0.65- 1.15)	0.90 (0.67- 1.20)	0.85 (0.63- 1.15)	0.91 (0.67-1.24)
Alcohol consumption				
Non	1.0 (reference)	1.0 (reference)	1.0 (reference)	1.0 (reference)
Does not drink every_day	0.96 (0.82- 1.13)	0.96 (0.81- 1.13)	0.97 (0.82- 1.15)	0.96 (0.80-1.15)

Drinks every day <	0.98 (0.84	0.00 (0.84	1.02 (0.86	
35 g/day	1.15)	1.16)	1.20)	1.03 (0.87-1.22)
Drinks every_day > 35 g/day	0.86 (0.64- 1.17)	0.90 (0.66- 1.21)	0.92 (0.68- 1.26)	0.91 (0.66-1.26)
Missing	1.14 (0.78- 1.66)	1.14 (0.78- 1.67)	1.24 (0.84- 1.83)	1.22 (0.82-1.83)
Random effects	,	,	e e e e e e e e e e e e e e e e e e e	
Community level variance (SE)	0.06 (0.05)	0.04 (0.06)	0.04 (0.07)	0.09 (0.04)

*HR for one point increment of community social support (range:

0-100)

for perteries only

STROBE Statement-checklist of items that should be included in reports of observational studies

	Item No	Recommendation	Page No
Title and abstract	1	(<i>a</i>) Indicate the study's design with a commonly used term in the title or	1
		the abstract	
		(b) Provide in the abstract an informative and balanced summary of what	2
		was done and what was found	
Introduction			
Background/rationale	2	Explain the scientific background and rationale for the investigation being reported	4-5
Objectives	3	State specific objectives, including any prespecified hypotheses	5
Methods			•
Study design	4	Present key elements of study design early in the paper	5
Setting	5	Describe the setting, locations, and relevant dates, including periods of	5-6
	C C	recruitment, exposure, follow-up, and data collection	
Participants	6	(a) Cohort study—Give the eligibility criteria and the sources and	5-6
1 uniterpunto	0	methods of selection of participants. Describe methods of follow-up	
		<i>Case-control study</i> —Give the eligibility criteria and the sources and	
		methods of case ascertainment and control selection. Give the rationale	
		for the choice of cases and controls	
		<i>Cross-sectional study</i> —Give the eligibility criteria, and the sources and	
		methods of selection of participants	
		(b) Cohort study—For matched studies, give matching criteria and	5-6
		number of exposed and unexposed	
		<i>Case-control study</i> —For matched studies give matching criteria and the	
		number of controls per case	
Variables	7	Clearly define all outcomes exposures predictors potential confounders	6-7
v unuoies	,	and effect modifiers. Give diagnostic criteria, if applicable	
Data sources/	8*	For each variable of interest give sources of data and details of methods	6-9
measurement	0	of assessment (measurement) Describe comparability of assessment	
		methods if there is more than one group	
Bias	9	Describe any efforts to address potential sources of bias	5-6
Study size	10	Explain how the study size was arrived at	5-6
Quantitative variables	11	Explain how quantitative variables were handled in the analyses. If	6-9
Qualificative variables		applicable describe which groupings were chosen and why	
Statistical methods	12	(a) Describe all statistical methods, including those used to control for	6-10
Statistical methods	12	confounding	0 10
		(b) Describe any methods used to examine subgroups and interactions	6-10
		(c) Explain how missing data were addressed	6-10
		(d) Cohort study. If applicable, explain how loss to follow up was	6
		addressed	0
		autrosou	
		controls was addressed	
		Cross-sactional study. If applicable describe analytical methods taking	
		cross-sectional study—in applicable, describe analytical methods taking	
		(a) Describe any consitiuity analyses	10
		(e) Describe any sensitivity analyses	10

Continued on next page

Results			
Participants	13*	(a) Report numbers of individuals at each stage of study—eg numbers potentially	5-0
		eligible, examined for eligibility, confirmed eligible, included in the study,	
		completing follow-up, and analysed	
		(b) Give reasons for non-participation at each stage	
		(c) Consider use of a flow diagram	
Descriptive	14*	(a) Give characteristics of study participants (eg demographic, clinical, social) and	10
data		information on exposures and potential confounders	
		(b) Indicate number of participants with missing data for each variable of interest	
		(c) Cohort study—Summarise follow-up time (eg, average and total amount)	6
Outcome data	15*	Cohort study—Report numbers of outcome events or summary measures over time	6
		Case-control study-Report numbers in each exposure category, or summary	
		measures of exposure	
		Cross-sectional study—Report numbers of outcome events or summary measures	
Main results	16	(a) Give unadjusted estimates and, if applicable, confounder-adjusted estimates and	10
		their precision (eg, 95% confidence interval). Make clear which confounders were	12
		adjusted for and why they were included	
		(b) Report category boundaries when continuous variables were categorized	10
			12
		(c) If relevant, consider translating estimates of relative risk into absolute risk for a	
		meaningful time period	
Other analyses	17	Report other analyses done-eg analyses of subgroups and interactions, and	11
		sensitivity analyses	12
Discussion			
Key results	18	Summarise key results with reference to study objectives	12
			1:
Limitations	19	Discuss limitations of the study, taking into account sources of potential bias or	14
		imprecision. Discuss both direction and magnitude of any potential bias	1:
Interpretation	20	Give a cautious overall interpretation of results considering objectives, limitations,	14
		multiplicity of analyses, results from similar studies, and other relevant evidence	15
Generalisability	21	Discuss the generalisability (external validity) of the study results	12
			13
Other informati	on		
Funding	22	Give the source of funding and the role of the funders for the present study and, if	10

*Give information separately for cases and controls in case-control studies and, if applicable, for exposed and unexposed groups in cohort and cross-sectional studies.

Note: An Explanation and Elaboration article discusses each checklist item and gives methodological background and published examples of transparent reporting. The STROBE checklist is best used in conjunction with this article (freely available on the Web sites of PLoS Medicine at http://www.plosmedicine.org/, Annals of Internal Medicine at http://www.annals.org/, and Epidemiology at http://www.epidem.com/). Information on the STROBE Initiative is available at www.strobe-statement.org.