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Association between change in handgrip strength and cognitive function in Korean adults: a longitudinal panel study

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Abstract

Background: Muscular function, such as handgrip strength, has been suggested as an associated factor for cognitive impairment. This study investigated the association between temporal change in handgrip strength and cognitive function using longitudinal, nationwide data from Korean older adults.

Methods: Our study used data from the Korean Longitudinal Study of Aging (KLoSA). The analysis covered 6696 participants who had taken the handgrip strength test and Mini-Mental State Examination (MMSE) from 2006 to 2018. We adopted general estimating equations to assess the temporal effect of handgrip strength change on cognitive function.

Results: After adjusting for covariates, we observed an association between handgrip strength and low MMSE scores ($\beta = -0.3142$ in men, $\beta = -0.2685$ in women). Handgrip strength as a continuous variable was positively correlated with MMSE scores after adjustment ($\beta = 0.0293$ in men, $\beta = 0.0347$ in women). The group with decreased handgrip strength over time also showed greater odds for mild cognitive impairment (OR = 1.23, 95%CI = 1.05–1.27 in men, OR = 1.15, 95%CI = 1.05–1.27 in women) and dementia (OR = 1.393, 95%CI = 1.18–1.65 in men, OR = 1.19, 95%CI = 1.08–1.32 in women).

Conclusions: This study identified the relationship between handgrip strength change and cognitive function among South Korean adults. According to our large, longitudinal sample, decreasing handgrip strength was associated with decline in cognitive function.

Keywords: Handgrip strength, Cognitive impairment, Cognitive function, Aging, Korean longitudinal study of aging

Background

As aging progresses, impairment in cognitive function may arise, and its intensified form, dementia, is considered a major health problem worldwide [1]. Cognitive impairment induces socio-economic burden by causing poor quality of life, hospitalization, increased mortality,

and poverty [2–5]. However, cognitive decline generally occurs gradually. Its onset or early phase is not easy to detect, with patients remaining undiagnosed until they display some functional impairment. Progression is likewise difficult to stop because cognitive impairment is a degenerative disease. Therefore, many studies have focused on methods to prevent cognitive impairment, including early detection and intervention, for example, by prescribing drugs such as donepezil, which is now widely used in clinical settings [6, 7].

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Studies have suggested the various factors associated with cognitive impairment and the ways to prevent decline based on those factors. The patient's neuropsychiatric condition, including depression, insomnia, or drug use [8–13], or chronic disease, such as hypertension or diabetes [14, 15], have been associated with cognitive decline. Other studies have pointed to several modifiable health behaviors including exercise, and body conditions, such as body mass index (BMI) and muscle mass, as being associated with dementia [16]. Among these, the relationship between handgrip strength, which can be easily measured, and cognitive impairment is becoming increasingly apparent [17].

A longitudinal study conducted in the United States positively linked handgrip strength with cognitive function, and a cohort study on Mexican Americans showed the association between baseline handgrip strength and Mini-Mental State Examination (MMSE) scores [18, 19]. A recent study using a longitudinal panel also showed that lower handgrip strength was associated with a higher odds ratio for cognitive impairment in aging Americans [20]. Similarly, a longitudinal study conducted in South Korea indicated the association between greater handgrip strength and a lower odds ratio of cognitive impairment [13]. However, few studies have investigated the association between changes in handgrip strength and cognitive impairment [17]. The confirmation of such an association in a large sample and through longitudinal design study would be useful as a basis for preventing cognitive decline by modulating handgrip strength via strength exercises. Therefore, we aimed to investigate the association between changes in handgrip strength and cognitive function in the Korean adult population based on a panel study, after adjusting for covariates that were assumed to affect cognitive function.

Methods

Study population and data

The data analyzed in this study were taken from the Korean Longitudinal Study of Aging database (KLoSA). The KLoSA is a longitudinal panel survey of nationally representative samples of community-dwelling adults aged above 45 years, and has been conducted every two years since 2006 [21]. The baseline data, gathered in 2006, include 10,254 Korean adults who have been interviewed by trained interviewers. The survey gathers information on respondents' family background, demographics, family composition, health, employment, income, assets, subjective expectations, and subjective quality of life. The seventh wave of KLoSA, conducted in 2018, covered an effective sample of 6136 from the original panels and 804 newly included panels. In the present study, we

employed survey data from 2006 to 2018, for a total of seven datasets. After deleting data with missing values for variables, we analyzed data from 6696 participants, including 2999 men and 3697 women. For statistical analysis, each change in handgrip strength from 2006 to 2018 was treated as an individual case rather than the population number itself. As KLoSA provides data in de-identified form which is open data for academic use, the need for informed consent was waived by the Institutional Review Board of Yonsei University's Health System (4–2021-0307).

Measures

Mini-mental state examination

To measure the cognitive function of the participants, we referred to their MMSE scores. The MMSE is a widely used tool for measuring cognitive function and screening for cognitive impairment in older adults [22, 23]. The validity of the Korean version of the MMSE (k-MMSE) has been established for its usefulness in screening for cognitive impairment [24]. With a total score of 30, the MMSE's cut-off level for mild cognitive impairment is 23 and that for dementia is 19 [25]. We used total scores for analysis to reveal detailed results regarding the association.

Handgrip strength

The KLoSA measures handgrip strength in kilograms using a handgrip dynamometer (Hand Grip Meter 6103, Tanita, Tokyo, Japan). Participants are asked to squeeze the dynamometer twice for each hand, and the mean value among four trials is recorded. Our data analysis excluded participants who declined to perform the test owing to physical problems. To analyze the association between changes in handgrip strength and cognitive function, we calculated the differences in reported values per wave. Handgrip strength was considered as both a categorized and a continuous variable in the analysis. As handgrip strength has been found to be significantly different between men and women in previous studies, we analyzed the data with stratification by sex [26, 27]. We also standardized the handgrip strength change by calculating the percentage change in handgrip strength from the handgrip strength of the original wave. The asymmetry in handgrip strength was calculated by subtracting the lower value from the higher value of handgrip strength. The change of the asymmetry per wave was then divided into two groups: 1) decreased, 2) same or increased.

Covariates

We considered demographic and health-related factors as covariates in the analysis. Demographic characteristics

included age, educational level, residential region, working status, household income, participation in social activities, and number of cohabiting generations. Health-related factors included smoking/alcohol use status, number of chronic medical conditions, BMI, and perceived health status. All the multivariable models controlled for all covariates unless stated otherwise.

Statistical analysis

All statistical analyses were performed separately for men and women to rule out the effect of sex in terms of the difference in handgrip strength on cognitive function. We employed analysis of variance to investigate and compare the general characteristics of the study population. We also constructed a generalized estimating equation model for regression analysis between MMSE scores and change in handgrip and other covariates. The analysis was conducted twice using the different variable types of change in handgrip strength: the two categorical groups of change in handgrip strength and the continuous variable of the same. The results were presented as regression coefficients (β) and 95% confidence intervals (95%CI). We performed subgroup analyses for a detailed study of the interaction between change in handgrip strength and other variables associated with MMSE scores. All analyses were carried out using SAS Version 9.4 (SAS Institute, Cary, North Carolina, USA). The results were considered statistically significant at $p < 0.05$.

Results

Table 1 gives the baseline characteristics of the study population stratified by sex. The unadjusted comparison showed no statistical difference in MMSE scores between the two groups of handgrip strength change in both sexes. Other covariates, such as age, educational level, region of residence, working status, household income, participation in social activities, BMI, and perceived health status, showed significant differences in MMSE scores for both sexes. The comparison of mean MMSE scores between two groups in Wave 2 and Wave 7 are presented in Supplementary Table 1. Both groups showed no statistically significant differences in mean MMSE scores for both Waves 2 and 7.

Table 2 shows the multiple regression analysis results for associations between MMSE scores and handgrip strength change groups after adjusting for covariates. Compared with the same or increased handgrip strength group, the decreased handgrip strength group showed highly significant regression coefficients (-0.3142 in men and -0.2685 in women). Decreased handgrip strength was associated with decreased MMSE scores in

both sexes. The results of other covariates are also shown in Table 2. Higher age showed a significant association with decreased MMSE scores, albeit with smaller regression coefficients. The decreased handgrip strength group also showed statistically significant odds ratios for mild cognitive impairment (OR = 1.23 in men, OR = 1.15 in women) and dementia (OR = 1.39 in men, OR = 1.19 in women), as illustrated in Table 3.

Table 4 shows the results of the multiple regression analysis between MMSE scores and handgrip strength change in continuous values with the same covariates in Table 2. The regression coefficients were 0.0293 in men and 0.0347 in women, indicating a high level of statistical significance. The results affirm the positive association between change in handgrip strength and MMSE total scores in both sexes. The decrease in handgrip strength asymmetry was associated with lower MMSE scores ($\beta = -0.1476$ in men and $\beta = -0.1755$ in women) as shown in Table 5.

Discussion

In this study, we identified that change in handgrip strength is associated with cognitive function in community-dwelling South Korean adults. The group with decreased handgrip strength was associated with low cognitive function when compared with the group with the same or increased handgrip strength. Furthermore, the value of the handgrip strength change was positively correlated with the MMSE scores in the study population.

Our results were generally consistent with previous studies on the relationship between change in handgrip strength and cognitive function in different populations. Previous studies showed that low baseline handgrip strength was associated with cognitive decline [28]. Christensen et al. reported that handgrip strength change, rather than initial strength, predicts changes in memory task performance [29]. MacDonald et al. suggested that biological changes, including grip strength change, share significant time-varying associations with change in cognitive function; handgrip strength decline is associated with cognitive function decline [29]. Several studies reported that physical frailty, including grip strength, was associated with cognition, suggesting that they might share common pathology [30, 31]. Compared with previous works, our results newly established that handgrip strength change is associated with cognitive function in older adults.

In our study, a decreased handgrip strength asymmetry was associated with a lower MMSE score, as shown in Table 5. McGrath et al. showed that handgrip strength

Table 1 Baseline characteristics of the study population according to Mini-Mental State Examination scores

	Men (N = 2999)				Women (N = 3697)				p-value	
	Participants		MMSE		Participants		MMSE			
	N	%	Mean ± S.D.	p-value	N	%	Mean ± S.D.	p-value		
Change in handgrip strength									0.6874	0.2907
Same or increased	1196	39.9	26.891	3.397	1673	45.3	25.385	4.798		
Decreased	1803	60.1	26.846	3.654	2024	54.7	25.253	4.813		
Age (years)									<0.0001	<0.0001
45–54	831	27.7	28.331	2.030	1131	30.6	27.842	2.396		
55–64	938	31.3	27.527	2.689	1084	29.3	26.522	3.404		
65–74	874	29.1	26.122	3.492	991	26.8	23.784	4.798		
≥75	356	11.9	23.517	5.396	491	13.3	19.904	6.070		
Education level									<0.0001	<0.0001
Elementary school or less	908	30.3	24.816	4.463	2016	54.5	23.321	5.293		
Middle school	522	17.4	27.080	2.905	635	17.2	27.047	2.981		
High school	1066	35.5	27.804	2.619	879	23.8	28.023	2.290		
University or beyond	503	16.8	28.346	2.095	167	4.5	28.503	1.951		
Region									<0.0001	<0.0001
Metropolitan	1240	41.3	27.432	3.045	1580	42.7	26.157	4.196		
Small or medium cities	991	33.0	26.898	3.499	1206	32.6	25.080	5.055		
Rural	768	25.6	25.904	4.138	911	24.6	24.157	5.176		
Working status									<0.0001	0.0198
Working	1905	63.5	27.569	2.685	1225	33.1	26.581	3.541		
Non-working	1094	36.5	25.637	4.438	2472	66.9	24.684	5.211		
Household income									<0.0001	0.0097
Quartile 1 (low)	600	20.0	24.760	4.534	991	26.8	23.188	5.256		
Quartile 2	816	27.2	26.799	3.210	987	26.7	25.401	4.440		
Quartile 3	846	28.2	27.508	2.959	907	24.5	26.114	4.484		
Quartile 4 (high)	737	24.6	27.910	2.837	812	22.0	26.905	4.043		
Participation in social activities									<0.0001	<0.0001
No	528	17.6	25.011	4.919	819	22.2	22.933	5.708		
Yes	2471	82.4	27.260	3.045	2878	77.8	25.990	4.283		
Smoking									0.0327	0.7566
Current	1179	39.3	27.160	3.145	107	2.9	24.019	4.939		
Former	778	25.9	26.719	3.533	38	1.0	24.395	5.274		
Never	1042	34.7	26.638	3.960	3552	96.1	25.362	4.792		
Alcohol Intake									0.1605	0.0340
Yes	1895	63.2	27.104	3.255	709	19.2	26.616	3.625		
No	1104	36.8	26.452	3.982	2988	80.8	25.004	4.997		
Number of chronic medical conditions									0.1283	0.5450
None	1539	51.3	27.402	3.153	1672	45.2	26.53	4.14		
1	895	29.8	26.535	3.730	1134	30.7	24.85	4.83		
≥2	565	18.8	25.920	4.010	891	24.1	23.62	5.31		
Number of cohabiting generations									0.0212	0.0541
Couple	1394	46.5	26.305	3.859	1783	48.2	24.812	4.731		
Two generations	1266	42.2	27.582	2.855	1415	38.3	26.339	4.443		
Over two generations	339	11.3	26.484	4.091	499	13.5	24.196	5.492		
BMI									0.0180	<0.0001
Underweight	89	3.0	24.944	4.971	114	3.1	23.360	6.044		
Normal weight	1321	44.0	26.513	3.782	1628	44.0	25.076	5.105		
Overweight	1012	33.7	27.292	3.290	1024	27.7	25.974	4.278		

Table 1 (continued)

	Men (N = 2999)				Women (N = 3697)			
	Participants		MMSE		Participants		MMSE	
	N	%	Mean ± S.D.	p-value	N	%	Mean ± S.D.	p-value
Obesity	544	18.1	27.270	2.917	818	22.1	25.571	4.389
Severe obesity	33	1.1	26.273	3.785	113	3.1	22.850	4.881
Perceived health status				<0.0001				<0.0001
Healthy	1752	58.4	27.744	2.675	1673	45.3	26.993	3.531
Average	855	28.5	26.035	3.803	1222	33.1	24.634	4.809
Unhealthy	392	13.1	24.742	4.866	802	21.7	22.843	5.710

BMI body mass index, Underweight: BMI < 18.5, Normal weight: 18.5 ≤ BMI < 23, Overweight: 23 ≤ BMI < 25, Obesity: 25 ≤ BMI < 30, Severe obesity: 30 ≤ BMI

asymmetry was associated with lower cognitive functioning [32]. The change in handgrip strength through aging or exercise could be different, resulting in a stronger or a weaker hand; thus, the change in asymmetry showed different patterns related to the asymmetry itself, with respect to association with cognitive function.

The etiology of the association between change in handgrip strength and cognitive function has not been established, although several possible explanations have been suggested. One is that physical activity increases the size of the prefrontal and hippocampal brain areas, thereby reducing cognitive decline [33–35]. Changes in handgrip strength can reflect the changes in the physical activity of individuals; thus, decreased handgrip might reflect reduced cognitive function. The frailty concept could be one explanation of the relationship between handgrip and reduced cognitive function. Handgrip strength decline could be an early and readily detectible indicator of frailty, which includes consequent decline of cognitive function in older adults. Previous studies showed that physical frailty, including being underweight, having weaker grip strength, and having a poor performance on the chair stand test was associated with cognitive decline [36, 37]. Another explanation is that cognitive function and handgrip strength might share a common domain of the brain, such as the frontal executive function area; decreased handgrip strength and cognitive decline might occur simultaneously.

Meanwhile, decreased cognitive function might also induce the change in handgrip strength. A study on the direction of the relationship between strength and cognitive function showed a significant bi-directional relationship [13]. Thus, muscular strength and cognitive function might share common causes of change. In our study, we could not exclude the possibility that participants with low cognitive function might have had difficulty in maintaining physical activity, including strength

exercises, which could lead to the decrease in their handgrip strength. Further research should be performed to test the directionality or causality of the two variables.

This study has several limitations. First, as the data were collected via a survey, the results might be biased. Second, we excluded the data of those who did not answer the important covariate questions, which may have induced the underestimation of cognitive decline in the participants. Third, we could not include biological risk factors, which could have led us to overlook some important confounding variables. Several biological factors have been established as risk factors of cognitive impairment in adults, and future studies should include these in regression model analyses [38]. Fourth, as we used brief measurements for cognitive function, the impact of handgrip strength changes on different neurocognitive domains could not be analyzed in this study. Previous studies showed that grip strength had different effects on cognitive domains [39, 40]. Further research using a comprehensive neurocognitive test would refine our study results. Finally, cause and effect could not be established because our study did not use a prospective design, which could be used to assess the causality of change in handgrip strength vis-à-vis change in cognitive function.

Nonetheless, the strengths of our study include the relatively large sample size and longitudinal design. Our results can be representative of the Korean adult population. Another strength is that this study used standardized tools to measure muscle strength and cognitive function; therefore, the results are readily applicable for further study. Moreover, given our use of the change in handgrip strength rather than baseline strength, the present results can be referenced when introducing lifestyle modifications, such as strength exercises, for older adults to help them maintain or increase their handgrip strength, which can prevent cognitive function decline.

Table 2 Results of the GEE analysis of handgrip strength change in two groups and Mini-Mental State Examination scores

	Men		Women	
	β	95%CI	β	95%CI
Changes in Handgrip strength				
Same or Increased	Ref.		Ref.	
Decreased	-0.3142	(- 0.4129–0.2154)	-0.2685	(- 0.3732–0.1637)
Age				
45–54	Ref.		Ref.	
55–64	-0.1218	(- 0.2560 0.0124)	-0.1374	(- 0.2760 0.0012)
65–74	-0.5936	(- 0.7915–0.3957)	-1.2507	(- 1.4799–1.0214)
≥ 75	-1.6447	(- 1.9444–1.3449)	-3.4250	(- 3.7685–3.0814)
Education level				
Elementary school or less	-2.2240	(- 2.4879–1.9601)	-2.9756	(- 3.2934–2.6577)
Middle school	-0.8777	(- 1.1216–0.6337)	-1.0498	(- 1.3567–0.7428)
High school	-0.5350	(- 0.7142–0.3557)	-0.5839	(- 0.8543–0.3136)
University or beyond	Ref.		Ref.	
Region				
Metropolitan	Ref.		Ref.	
Small or Medium Cities	-0.3659	(- 0.5544–0.1774)	-0.5500	(- 0.7737–0.3262)
Rural	-0.3564	(- 0.5755–0.1373)	-0.8134	(- 1.0690–0.5578)
Working status				
Working	Ref.		Ref.	
Non-working	-0.4528	(- 0.6053–0.3003)	-0.4327	(- 0.5817–0.2836)
Household income				
Quartile 1 (low)	-0.5991	(- 0.8564–0.3417)	-0.3133	(- 0.5657–0.0609)
Quartile 2	-0.0653	(- 0.2554 0.1249)	0.0357	(- 0.1743 0.2458)
Quartile 3	-0.0110	(- 0.1703 0.1484)	-0.0498	(- 0.2293 0.1296)
Quartile 4 (high)	Ref.		Ref.	
Participation in social activities				
No	-0.9662	(- 1.1680–0.7644)	-1.2188	(- 1.4130–1.0247)
Yes	Ref.		Ref.	
Smoking				
Current	0.2185	(- 0.1002 0.3107)	0.2520	(- 0.3558 1.0495)
Former	0.1053	(0.0150 0.4221)	0.3468	(- 0.3385 0.8425)
Never	Ref.		Ref.	
Alcohol Intake				
Yes	0.0944	(- 0.0633 0.2521)	0.3999	(0.2034 0.5964)
No	Ref.		Ref.	
Number of chronic medical conditions				
None	Ref.		Ref.	
1	-0.0207	(- 0.1878 0.1464)	0.0023	(- 0.1920 0.1966)
≥ 2	-0.1014	(- 0.3174 0.1145)	-0.2783	(- 0.5405–0.0161)
Number of cohabiting generations				
Couple	0.4467	(0.1589 0.7346)	0.7265	(0.4234 1.0296)
Two generation	0.3116	(0.0398 0.5835)	0.4248	(0.1206 0.7291)
Over two generation	Ref.		Ref.	
BMI				
Underweight	0.2462	(- 0.2392 0.7316)	-0.3675	(- 0.8535 0.1185)
Normal weight	Ref.		Ref.	
Overweight	0.1452	(- 0.0042 0.2945)	0.3958	(0.2294 0.5622)
Obesity	0.1092	(- 0.0672 0.2856)	0.5409	(0.3336 0.7483)
Severe obesity	0.2504	(- 0.4315 0.9323)	-0.2268	(- 0.7595 0.3060)
Perceived health status				
Healthy	Ref.		Ref.	
Average	-0.2811	(- 0.4087–0.1536)	-0.3218	(- 0.4640–0.1796)
Unhealthy	-1.2703	(- 1.5197–1.0208)	-1.1481	(- 1.3674–0.9288)

BMI body mass index, Underweight: BMI < 18.5, Normal weight: 18.5 \leq BMI < 23, Overweight: 23 \leq BMI < 25, Obesity: 25 \leq BMI < 30, Severe obesity: 30 \leq BMI

Table 3 Results of the GEE analysis of handgrip strength change and mild cognitive impairment/dementia

	Mild Cognitive Impairment (20 ≤ MMSE < 24)				Dementia (MMSE < 20)			
	Men		Women		Men		Women	
	Adjusted OR	95%CI	Adjusted OR	95%CI	Adjusted OR	95%CI	Adjusted OR	95%CI
Changes in Handgrip strength								
Same or Increased	Ref.				Ref.			
Decreased	1.23	(1.07–1.41)	1.15	(1.05–1.27)	1.39	(1.18–1.65)	1.19	(1.08–1.32)

*All variables in Table 2 were included in the GEE model

Table 4 Results of the GEE analysis of handgrip strength change in continuous variables and Mini-Mental State Examination scores

	Men		Women	
	β	95%CI	β	95%CI
Handgrip strength change (kg)	0.0293	(0.0210 0.0375)	0.0347	(0.0232 0.0463)
Handgrip strength change (%)	0.0077	(0.0052 0.0101)	0.0049	(0.0030 0.0068)

*All variables in Table 2 were included in the GEE model. Handgrip strength change were shown in absolute value (kg) and percent change from previous handgrip strength (%)

In conclusion, this study identified the relationship between changes in handgrip strength and cognitive function among South Korean adults. Decreased handgrip strength was associated with cognitive decline in our longitudinal, large-sample study. Further studies exploring the underlying mechanisms of the association between handgrip strength and cognitive impairment, as well as the preventive effect of increasing the former, could provide valuable strategies for treating and preventing cognitive impairment in clinical settings.

Table 5 Results of the GEE analysis of handgrip strength asymmetry change and Mini-Mental State Examination scores

	Men		Women	
	β	95%CI	β	95%CI
Handgrip strength asymmetry change (kg)				
Same or Increased	Ref.		Ref.	
Decreased	-0.1476	(-0.2485–0.0466)	-0.1755	(-0.2822–0.0688)

*All variables in Table 2 were included in the GEE model. Handgrip strength asymmetry was calculated by subtracting lower handgrip strength from higher handgrip strength. The change of handgrip strength asymmetry between two years were grouped into Same/Increased and Decreased

Supplementary Information

The online version contains supplementary material available at <https://doi.org/10.1186/s12877-021-02610-2>.

Additional file 1: Supplementary Table 1. MMSE scores of Same or Increased and Decreased handgrip strength group in Wave 2 and Wave 7.

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Authors' contributions
HK, SHK and WJ had full access to all the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis. Concept and design: HK, YK. Acquisition, analysis, or interpretation of data: HK, WJ. Drafting of the manuscript: HK. Critical revision of the manuscript for important intellectual content: ECP, SIJ, YK. Statistical analysis: HK, WJ, SHK. Supervision: YK. All authors read and approved the manuscript.

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Availability of data and materials
The datasets analyzed during the current study are available in the KLoSA repository, <https://survey.keis.or.kr/eng/klosa/databoard/List.jsp>.

Declarations

Ethics approval and consent to participate
The KLoSA study was approved by the Statistics Korea of Korean Government (Approval number: 33602) and Institutional Review Board of Korea National Institute for Ethics Policy (P01–201909–22-002). The survey was conducted after acquiring verbal consent of the participants by the trained study interviewer. This study was exempted from approval by the Institutional Review Board of Yonsei University's Health System (4–2021-0307), adhering to the principles of the Declaration of Helsinki.

Consent for publication
Not applicable.

Competing interests
The authors declare that they have no competing interests.

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