Association of Frailty Status with Risk of Fall among Middle-Aged and Older Adults in China: A Nationally Representative Cohort Study

Z. Lu¹, Y. Er¹, Y. Zhan^{2,3}, X. Deng¹, Y. Jin¹, P. Ye^{1,4}, L. Duan¹

1. National Center for Chronic and Noncommunicable Disease Control and Prevention, Chinese Center for Disease Control and Prevention, Beijing, China; 2. Department of Epidemiology and Biostatistics, School of Population Medicine and Public Health, Chinese Academy of Medical Sciences and Peking Union Medical College, Beijing, China; 3. School of Public Health, LKS Faculty of Medicine, The University of Hong Kong, Hong Kong Special Administrative Region, China; 4. School of Population Health, Faculty of Medicine and Health, University of New South Wales, Sydney, Australia.

Corresponding Author: Leilei Duan and Pengpeng Ye National Center for Chronic and Noncommunicable Disease Control and Prevention, Chinese Center for Disease Control and Prevention, Beijing, China, E-mail: duanleilei@ncncd,chinacdc.cn, yepengpeng@ncncd,chinacdc.cn

Abstract

OBJECTIVES: To investigate the association between frailty status and risk of fall among middle-aged and older Chinese people.

DESIGN: A nationally representative prospective cohort study.

SETTING AND PARTICIPANTS: The study included 13,877 community-dwelling participants aged 45 years and above from the China Health and Retirement Longitudinal Study.

METHODS: Frailty status was identified by the frailty index of health deficit accumulation. 34 variables at baseline were selected to calculate the frailty index. We excluded participants with incomplete information in construction of the frailty index at baseline. Falls were measured based on the respondents' self-report. We used a logistic regression model to estimate the associations between the frailty status and risk of fall, and subgroup analyses and sensitivity analyses were further conducted.

RESULTS: Of 13,877 participants, 2310 (16.6%) had falls during the observation period. 9027 (65.0%) participants were classified as robust, 4019 (29.0%) participants were classified as pre-frail, and 831 (6.0%) participants were classified as frail. Our results indicated per 0.01 increment in the frailty index was significantly associated with an increased risk of fall among middle-aged and older participants (OR: 1.52, 95%CI: 1.45-1.60). Such association was stronger when frailty was presented as a categorical variable, with an OR of 1.75 (95%CI: 1.59-1.93) for pre-frail and 3.04 (95%CI: 2.60-3.56) for frail. The area under the curve of the logistic model was 0.612 (95%CI: 0.600-0.625). Each 0.01 increment of the frailty index was association with a higher risk of fall among middle-aged (45-59years) participants (OR: 1.28, 95%CI: 1.26-1.41) at baseline (p=0.015 for interaction).

CONCLUSION: Frailty was significantly associated with an increased risk of fall among community-dwelling middle-aged and older people in China. It is necessary to screen and recognize frailty status to prevent falls among middle-aged and older adults.

Key words: Frailty, deficit accumulation, falls, cohort study.

Introduction

ith the population ageing rapidly, falls have become one of the significant public health problem worldwide (1, 2). In 2019, 703 million people reached 65 years old or above in the world, and an estimated number of older people is projected to 1.5 billion in 2050 (3). Approximately one-third community-dwelling adults aged 65 years and above and one-half of adults older than 85 years of age are reported falls each year (4). Globally, it was estimated falls as the second leading cause of death from unintentional injuries after road traffic injuries (5). In addition to death, falls can result in limited mobility, fractures, disabilities, and hospitalization (6, 7). Moreover, falls are likely to compromise quality of life and cause large consumption of medical resources (8, 9). In China, falls are the leading cause of injury death in the elderly (10). According to the Global Burden of Disease Study, the incidence, mortality, and disabilityadjusted-life-years for falls among Chinese people increased from 1990 to 2017 (11). Falls have a huge impact on the burden of disease among the middle-aged population, which are the second leading cause of injury deaths among middle-aged people. In addition, a majority of studies on the associations between frailty index and risk of fall were conducted among older people (over 60 years), resulting in limited evidence regarding the middle-aged population. Therefore, it is necessary to investigate the risk factors of falls in the middle-aged and older people.

Frailty is an age-related biological syndrome characterized by a reduction of physiologic reserve and an increase of vulnerability to stressors, which is strongly associated with adverse health outcomes, such as disability, delirium, and hospitalization (12). With the acceleration of aging, frailty, as a comprehensive index to measure the physiology, function and psychology of the elderly, has been as focal points of academic circles. A previous systematic review reported the prevalence of frailty in community-dwelling adults aged 65 years and above was ranging from 4.0% to 59.1% (13). Frailty index is the most widely used method for assessing frailty status, which is also called deficits accumulation index (14).

Previous studies showed an association of frailty with incidence of falls in the elderly (15, 16). However, most of these studies were carried out in developed countries. Studies on the association between frailty and falls are scanty in China, and most of them were limited to single-centered (17) and cross-sectional design (18). There is a lack of nationally representative longitudinal studies. Thus, in this study, we aim

to investigate the association between frailty index and risk of fall among middle-aged and older adults by using data of the China Health and Retirement Longitudinal Study (CHARLS).

Methods

Study design and Sampling

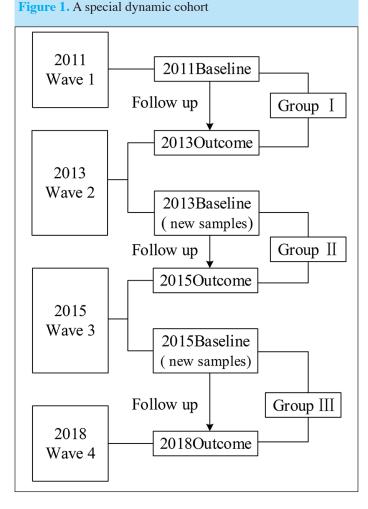
The study data was obtained from the CHARLS, an ongoing nationally representative cohort study of the residents in mainland China aged 45 years and above. Further details of the study design have been described elsewhere (19). Briefly, a total of 17,708 respondents in 10,257 households were chosen in 28 provinces, 150 country-level units and 450 urban communities/villages, using a multi-stage stratified probability-proportional-to-size random sampling strategy. The baseline survey was conducted in June 2011 with a response rate of 80.5%, and follow-up surveys were carried out every two years subsequently (i.e. wave 2 in 2013, wave 3 in 2015 and wave 4 in 2018, respectively). Face-to-face computer assisted personal interview was adopted by trained staff using a standardized questionnaire on sociodemographic characteristics, lifestyle factors, mental and physical health status, and physical measurements. The CHARLS study had been reviewed and approved by the Institutional Review Board (IRB) at Peking University.

As the study progressed, new samples were added to the study in each wave to ensure the sample fully represent those aged 45 years and above. The CHARLS baseline data was used to calculate the frailty index while fall information was obtained from the CHARLS follow-up data. Because the dates of falls were not available, analyses were performed only using the data for falls during the two or three years of followup. We constructed a special dynamic cohort, including three groups. In group I, baseline data were from wave 1 (2011), and fall information was from wave 2 (2013); in group II, new participants were added from the refresher sample in wave 2 (2013), making wave 2 as their baseline data, and fall information was from wave 3 (2015); and in group III, new participants were added from the refresher sample in wave 3 (2015), making wave 3 as their baseline data, and fall information was from wave 4 (2018) (Figure 1).

Measures

Frailty

Frailty status was measured by a frailty index, which was calculated by baseline deficits accumulation in health of each participant. The frailty index was constructed following a standard procedure (20). These deficits associated with health status included the following basic criteria: 1) The deficit covers a range of body systems; 2) the prevalence of the deficit generally increases with age; 3) the baseline prevalence of the deficit is more than 0.5% in the CHARLS study; and 4) the deficit does not saturate too early.



According to the CHARLS data and previous studies (21-23), we selected 34 variables at baseline to calculate the frailty index covering symptoms, body functions, disabilities, medical condition, and activities of daily living (Supplementary Table 1 in the Appendix). The deficit was dichotomized into 0 and 1 if the deficit was binary variables, with '0' referring to no deficit and '1' referring to the maximum deficit. For activities of daily living variable, the deficit was mapped into the 0-1 interval, where we assigned a 0 for the answer 'have no difficulty', 0.33 for 'have difficulty but can still do it', 0.67 for 'have difficulty and need help', and 1 for 'cannot do it'. For each participant, the frailty index was calculated by dividing the total number of the deficit in a person by 34 deficits considered. For example, if six deficits (each of four deficits scored 1 point and each of two deficits scored 0.67 point) were present in a participant, the frailty index would be (4x1+2x0.67)/34=0.16. Frailty states were divided into three level with the reference of previous study (21, 24, 25): frailty index ≤ 0.10 denoted the robust, frailty index >0.1 to <0.25 denoted pre-frail, and frailty index \geq 0.25 denotes frail.

Fall

The outcome of interest in the present study was fall in Chinese adults aged 45 years and above, which was assessed by the following single question: 'have you fallen down in the last two years?' Participants were asked to answer 'Yes' or 'No' (coding 1 or 0).

Covariates

Sociodemographic information, lifestyle and health related factors were collected at baseline. Age was categorized as 45-59 and \geq 60 years. Educational level was divided into four groups: no formal education, elementary school, middle or high school, and college or above. Marital status was classified as married and other status (divorced, widowed, and single). Lifestyle included smoking (current, past, or never) and drinking status (>1 per month, \leq 1 per month, or no). Health related factors included comorbidity, history of medication use (hypertension medications, diabetes medications, psychotropic medications), history of falls, eyesight, hearing, balanced performance (semitandem stand and full-tandem stand) and body mass index (BMI). BMI was calculated as weight in kilograms divided by the square of height in meters.

Statistical analysis

Data were presented as means (standard deviation, SD) for normally distributed continuous variables, and as medians (interquartile range, IQR) for non-normally distributed continuous variables. Categorical variables were presented as frequency with percentage (%). We described and compared the baseline characteristics of participants according to three categories of frailty status (robust, pre-frail, and frail) using the Chi-square test for categorical variables and analysis of variance or the Kruskal-Wallis test for continuous variables where appropriate. We excluded participants with incomplete information in construction of the frailty index and younger than 45 years at baseline. For missing data, it was assumed that the data were missing at random, and then, multiple imputation of chained equations strategy was performed by using other baseline variates. Ten imputed datasets were created and the results were pooled using Stata software version 14.0 command "mi estimate". To examine the association between frailty index and risk of fall, a series of logistic regression models were applied to calculated the odds ratio (OR) with 95% confidence interval (95% CI), with or without adjustment. Three models were built: in model 1, only frailty was included; in model 2, age and gender were adjusted; in model 3, the variables in model 2 plus residence, educational level, marital status, smoking, alcohol use, comorbidity, history of medication use, history of falls, eyesight, hearing, BMI, balance performance, follow-up period and group were adjusted. The receiver operating characteristic (ROC) curve was constructed to calculated the area under the curve (AUC) in order to evaluate frailty index's ability to predict risk of fall.

Subgroup analyses were conducted to investigate whether the potential association between frailty and risk of fall was affected by age (middle-aged: 45-59 vs. older: ≥ 60 years), gender (male vs. female), residence (rural vs. urban), marital status (married vs. others), history of falls (yes vs. no), alcohol use (>1 per month vs. \leq 1 per month vs. don't use), hypertension medications use (yes vs. no), diabetes medications use (yes vs. no). We used interaction terms (frailty was regarded as a continuous variable) and generalized hausman test (frailty was regarded as a categorical variable) to evaluate p values for interaction. Sensitivity analyses of different cutoff points of the frailty index (robust: \leq 0.1; pre-frail: 0.1-0.21; frail: \geq 0.21) were also conducted to evaluate the association results between frailty status and risk of fall.

All statistical analyses were performed using Stata software version 14.0 (Stata Corp, College Station, Texas). P values were two-sided test, with α less than 0.05 to be considered statistically significant.

Results

Baseline characteristics of participants

The ROC curve showed a fair accuracy of frailty index predicted risk of fall with an AUC of 0.612 (95%CI: 0.600-0.625). Of the 13,877 middle-aged and older participants in the analysis, 12,109 from group I, 1528 from group II, and 240 from group III. 9027 (65.0%) adults were classified as robust, 4019 (29.0%) adults were classified as pre-frail, and 831 (6.0%) adults were classified as frail. The mean frailty index was 0.092±0.086. Their mean BMI was 23.51±3.63 kg/m². The mean age of participants at baseline was 58.71±9.41 years. About three fifths (59.1%) of participants were aged younger than 60 years, and about half (48.6%) of the participants were male. Table 1 illustrated baseline characteristics of participants according to frailty status in this study. Frail persons were more likely to be older, be female, have no formal education, live in rural areas, be unmarried, be former and never smokers, be nondrinkers, have comorbidity, use medication for hypertension, diabetes and psychiatric illnesses, have a history of falls, wear glasses or corrective lenses, have poor hearing, and bad balance performance (keep semi-tandem stand for less than 10s or keep full-tandem stand for less than 30s/60s) at baseline.

Frailty status and risk of fall

Of the 13,877 included participants, 2310 (16.6%) had falls during the observation period. The incidence rate of falls was 13.3% among robust participants, 21.1% among pre-frail participants and 31.8% among frail participants. Table 2 showed that frailty status was significantly associated with risk of fall in all models. In the univariate analysis (model 1), when frailty was regarded as a continuous variable, the OR for risk of fall was 1.52 (95%CI: 1.45-1.60) per 0.01 increment of the frailty index. When frailty was regarded as a categorical variable (frail, pre-frail, and robust), compared with robust group, pre-frail group had a higher risk of fall (OR: 1.75, 95%CI: 1.59-1.93), similar as frail group (OR: 3.04, 95%CI: 2.60-3.56). In the multivariate analysis (model 3), the OR for risk of fall was 1.35 (95%CI: 1.26-1.45) per 0.01 increase of the frailty index. Compared with robust group, pre-frail group and frail group

Characteristics	cs of Participants Accord Total n(%) (n=13877)			P Value	
		Robust (n=9027)	Frailty status n(%) Pre-frail (n=4019)	Frail (n=831)	
Age					< 0.001
45-59 years	7690(59.1)	5583(66.0)	1850(48.8)	257(33.3)	
≥60 years	5330(40.9)	2876(34.0)	1939(51.2)	515(66.7)	
Gender					<0.001
Male	6738(48.6)	4767(52.9)	1666(41.5)	305(36.7)	
Female	7124(51.4)	4251(47.1)	2348(58.5)	525(63.3)	
Place of residence					< 0.001
Rural area	10843(78.2)	6923(76.7)	3245(80.8)	675(81.4)	
Urban area	3025(21.8)	2098(23.3)	773(19.2)	154(18.6)	
Educational level					< 0.001
No formal education	3546(25.6)	1897(21.0)	1280(31.9)	369(44.4)	
Elementary school	5607(40.4)	3510(38.9)	1763(43.9)	334(40.2)	
Middle or high school	4385(31.6)	3348(37.1)	914(22.7)	123(14.8)	
College or above	335(2.4)	269(3.0)	61(1.5)	5(0.6)	
Marital status	555(2.1)	207(0.0)	01(1.5)	5(0.0)	< 0.001
Married	12341(89.0)	8228(91.2)	3441(85.6)	672(81.0)	\$0.001
Divorced/widowed/single	1530(11.0)	794(8.8)	578(14.4)	158(19.0)	
Smoking	1550(11.0)	794(0.0)	576(14.4)	156(19.0)	< 0.001
Current	4287(30.9)	2025(22.6)	1060(26.6)	182(22.0)	<0.001
		3035(33.6)	1069(26.6)	183(22.0)	
Past	1226(8.8)	704(7.8)	413(10.3)	109(13.1)	
Never	8360(60.3)	5285(58.6)	2536(63.1)	539(64.9)	0.001
Alcohol use		2(25(20.2)	0(0(01.4)	100/15 5	<0.001
>1 per month	3624(26.1)	2635(29.2)	860(21.4)	129(15.5)	
≤1 per month	1094(7.9)	762(8.4)	296(7.4)	36(4.3)	
No	9157(66.0)	5628(62.4)	2863(71.2)	666(80.2)	
Comorbidity					<0.001
Yes	2550(18.4)	216(2.4)	1736(43.2)	589(72.0)	
No	11327(81.6)	8811(97.6)	2283(56.8)	233(28.0)	
History of medication use					
Hypertension medications, yes	2594(18.7)	1026(11.4)	1199(29.8)	369(44.4)	<0.001
Diabetes medications, yes	586(4.2)	178(2.0)	288(7.2)	120(14.4)	<0.001
Psychotropic medications, yes	62(0.5)	16(0.2)	28(0.7)	187(2.2)	<0.001
History of falls					< 0.001
Yes	2181(15.7)	970(10.8)	918(22.9)	293(35.3)	
No	11688(84.3)	8053(89.2)	3097(77.1)	538(64.7)	
Eyesight (wear glasses or corrective lenses)					<0.001
Yes	1817(13.1)	1086(12.0)	601(15.0)	130(15.7)	
No	12058(86.9)	7940(88.0)	3418(85.0)	700(84.3)	
Hearing					<0.001
Good or above	6350(45.7)	4858(53.8)	1327(33.0)	165(19.8)	
Fair	5740(41.4)	3452(38.3)	1921(47.8)	367(44.2)	
Poor	1786(12.9)	716(7.9)	771(19.2)	299(36.0)	
BMI, mean(SD)	23.51(3.63)	23.44(3.45)	23.61(3.94)	23.77(3.95)	<0.001
Balance performance					
Semi-tandem stand					<0.001
≥10s	11332(98.0)	7469(99.4)	3314(97.2)	549(86.3)	
<10s	229(2.0)	48(0.6)	94(2.8)	87(13.7)	
Full-tandem stand*					< 0.001
	9919(79.4)	6203(83.4)	2328(71.0)	287(53.8)	
≥30/60s	8818(78.4)	0205(05.4)	2520(71.0)	207(55.0)	

*For full-tandem stand, the 30s was for participants aged 70 years or older, and the 60s was for participants younger than 70 years.

Frail

Table 2. Association of Frailty and Risk of Fall								
	Falls		Model 1 *		Model 2 **		Model 3 ***	
	Cases, No	Incidence rate(%)	OR(95% CI)	P value	OR(95% CI)	P value	OR(95% CI)	P value
Continuous								
Frailty index (per 0.01 increase)	2310	16.6	1.52(1.45-1.60)	< 0.001	1.43(1.36-1.50)	<0.001	1.35(1.26-1.45)	< 0.001
Categorical								
Robust	1198	13.3	Ref.	Ref.	Ref.	Ref.	Ref.	Ref.
Pre-frail	848	21.1	1.75(1.59-1.93)	< 0.001	1.59(1.44-1.75)	< 0.001	1.37(1.21-1.55)	< 0.001

31.8

OR, odds ratio. 95% CI, 95% confidence interval; * Basic model: only frailty included in the model; ** Age-gender adjusted model: basic model + age, gender; *** Full adjusted model: age-gender adjusted model + residence, educational level, marital status, smoking, alcohol use, comorbidity, history of medication use, history of falls, eyesight, hearing, BMI, balance performance, follow up period and group.

3.04(2.60-3.56)

< 0.001

2.54(2.16 - 3.00)

Figure 2. Association between Frailty and Risk of Fall Stratified by Different Factors

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	A. Per-0.01 increase	B1. Pre-frail	B2. Frail
Subgroup	P for interaction	P for interaction	P for interaction
Age strara	0.015	0.142	0.701
45-59	+		— — —
≥60	+	-8-	_
Gender	0.381	0.118	0.163
Male	+		∎ →
Female	+		_
Residence	0.923	0.151	0.915
Rural	+		_
Urban			──■ ──→
Marital status	0.111	0.301	0.389
Married	-		_
Others			
History of falls	0.006	0.216	0.278
Yes	-		_
No	+		_
Alcohol use	0.464	0.277	0.128
>1 per month		_ 	_ >
≤1 per month		_	← ∎
No	+	-=-	_
Comorbidity	0.453	0.831	0.894
Yes	+		_
No	+		
Hypertension medications use	0.311	0.262	0.731
Yes		_ 	_
No	-	-8-	_
Diabetes medications use	0.442	0.741	0.951
Yes			
No	+		_
F			
0.5	1 1.5 2 2.5 3 OR (95% CI)	0.5 1 1.5 2 2.5 3 OR (95% CI)	0.5 1 1.5 2 2.5 3 OR (95% CI)

OR, odds ratio. 95% CI, 95% confidence interval; Model adjusted age, gender, residence, educational level, marital status, smoking, alcohol use, comorbidity, history of medication use, history of falls, eyesight, hearing, BMI, balance performance, follow up period and group. A: continuous variable. B1 and B2: categorical variable, taking robust as reference.

had a 37% and 97% higher risk of fall, respectively (OR: 1.37, 95%CI: 1.21-1.55 for pre-frail group; OR: 1.97, 95%CI: 1.59-2.44 for frail group).

Subgroup analyses

Figure 2 showed the results of the association between frailty status and risk of fall stratified by other potential risk factors. When frailty status was considered as a continuous variable (Figure 2-A), we found that the association was stronger among middle-aged participants (OR: 1.44, 95%CI: 1.29-1.60) than older participants (OR: 1.28, 95%CI: 1.16-1.41) at baseline (p=0.015 for interaction). Frailty status was associated with a

higher risk of falls in participants with no history of falls (OR: 1.41, 95%CI: 1.30-1.53), compared with those with a history of falls (OR: 1.23, 95%CI: 1.08-1.39) at baseline (p=0.006 for interaction). However, when frailty status was considered as a categorical variable (Figure 2-B), we could not see any statistically significant heterogeneity in all subgroups (p>0.05 for interaction).

1.97(1.59-2.44)

< 0.001

< 0.001

Sensitivity analyses

In sensitivity analyses, the results had marginal changes after replacing the cutoff point of frailty to 0.21. In the multivariate analysis, the OR for risk of fall was 1.35 (95%CI: 1.26-1.45)

Table 3. Sensitivity Analysis of Association between Frailty and Risk of Fall								
	Model 1 *		Model 2 **		Model 3 ***			
	OR(95% CI)	P value	OR(95% CI)	P value	OR(95% CI)	P value		
Continuous								
Frailty index(per 0.01 increase)	1.52(1.45-1.60)	< 0.001	1.43(1.36-1.50)	< 0.001	1.35(1.26-1.45)	<0.001		
Categorical								
Robust	Ref.	Ref.	Ref.	Ref.	Ref.	Ref.		
Pre-frail	1.67(1.51-1.85)	< 0.001	1.53(1.38-1.69)	< 0.001	1.34(1.18-1.51)	<0.001		
Frail	2.76(2.42-3.15)	< 0.001	2.34(2.04-2.68)	< 0.001	1.87(1.56-2.25)	<0.001		

OR, odds ratio. 95% CI, 95% confidence interval. * Basic model: only frailty included in the model. ** Age-gender adjusted model: basic model + age, gender. *** Full adjusted model: age-gender adjusted model + residence, educational level, marital status, smoking, alcohol use, comorbidity, history of medication use, history of falls, eyesight, hearing, BMI, balance performance, follow up period and group.

per 0.01 increase of the frailty index. Compared with robust group, both pre-frail group (OR: 1.34, 95%CI: 1.18-1.51) and frail group (OR: 1.87, 95%CI: 1.56-2.25) had higher risks of fall (Table 3).

Discussion

This study investigated the association between frailty status and risk of fall in a nationally representative longitudinal study of 13,877 participants aged 45 years and above. At baseline, the prevalence of frailty was 6.2% by using the measurement method of frailty index. At follow up, 16.5% of the participants experienced falls. The major finding of our study showed that there was a statistically significant association between frailty status and risk of fall among middle-aged and older adults in China. Each 0.01 increase of the frailty index was significantly associated with a higher risk of fall among middleaged participants (45-59 years) than older participants (60 years and older).

The frailty index was developed based on the theory of health deficit accumulation, combing a variety of complex health information into a single indicator, which could break the limitations of describing health status by using a single variable. The frailty index model was widely used in developed countries to evaluate the frailty status of the elderly, which has been proved to be reliable and valid (26). Furthermore, frailty index also had good repeatability. Although different countries used different data to evaluate the frailty status of the elderly by frailty index method, nearly all of the studies got the similar results (27). Previous studies also proved the validity of the frailty index model among adults aged 55 years and older in China (28). There is no international consensus for the healthrelated variables used to construct frailty index, although 30-40 variables included in an index have been proposed to accurately predict adverse outcomes (29). Hence we selected 34 variables to construct the frailty index. According to different levels of frailty, the general information of participants was analyzed. The results showed a higher level of frailty in female than in male, consistent with the findings of former studies (21, 30). Participants who had a history of falls were frailer than those who had not. For example, Fang et al (17) found that the frailty index of the elderly with recurrent falls was significantly higher than those without falls ($0.15\pm0.08 \text{ vs } 0.09\pm0.07$). The prevalence of frailty was 9.7% in adults aged 60 years and above. A study including seven cities reported similar results to our study, with the overall weighted frailty prevalence of 9.9% among Chinese population in the community aged 60 years and above (31).

In our study, logistic regression was used to analyze the relationship between frailty status and risk of fall. The results showed that frailty was associated with an increased risk of fall, even after adjusting for baseline covariates such as sociodemographic characteristics and lifestyle and health related factors. A Canadian study of nearly 4000 women aged 55 years and over, using 34 health deficits to construct a frailty index, found that per 0.01 increment in the frailty index was significantly associated with an increased risk of fall during short term of follow up (OR:1.02, 95%CI:1.02-1.03) (32), which showed similar results to our study. Moreover, a metaanalysis including 68,723 older individuals also found that frailty (OR:1.84, 95% CI:1.43-2.38) and pre-frailty (OR:1.25, 95%CI:1.01-1.53) were significantly associated with higher risks of fall, respectively (33). The results of the association between frailty index and risk of fall in subgroup analysis showed that frail middle-aged people aged 45-59 years were more vulnerable to fall, which suggested that the middleaged population should also be the targeted population for frailty intervention in addition to the elderly population. The prevention and treatment of frailty is still in the exploratory stage, and clinical trials of frailty intervention are sparse, such as medication management (34), exercise and nutrition support (35). However, early identification and intervention of frailty are important to prevent falls. Compared with other studies (22, 32), the present study used specific cutoff points to define frailty. Sensitivity analysis suggested that the relationship between frailty and risk of fall was not affected by slight fluctuations in cutoff points of frailty. However, the frailty cutoff point of Chinese population is still required to be validated in future research.

The mechanisms that underlie the association of frailty with increased risks of fall have been proposed. Balance and gait disorders are typical features of frailty and are major risk factors for falls (12). Moreover, frailty is often accompanied by muscle weakness, decreased bone mineral density, and chronic disease (36). All of these symptoms are likely to increase the risk of fall (37, 38).

There are several strengths in our study. First, this study was a nationally representative prospective cohort study of participants aged 45 years and over. Results in our study could be used as a reference to prevent falls among middle-aged and older people living in community in China. Second, the method of health deficit accumulation was used to calculate the frailty index, and continuous index can be used to analyze the trend association. Third, this study included a new sample from each follow-up, which increased the sample size and ensured the representativeness of the population aged 45 years and older. Considering the heterogeneity among samples in different periods, we adjusted the enrollment year and follow-up period in the analysis to reduce bias. Meanwhile, we acknowledge some limitations. First, similar to previous large-scale studies (32), the data for falls was collected by respondents' selfreport. However, the validity of self-reported fall in the CHARLS study has been confirmed only by examining the relationship of balanced performance and falls (39). Second, some confounding factors of the association of frailty with incident fall, such as physical environment, sarcopenia, and nutrients intake were not adjusted in our study. Because these variables were not available in CHARLS data. In addition, dates for falls and frequency of falls were also unavailable. Finally, the response rate is 80.5% at baseline, which may have slight impact on our findings. However, CHARLS, a first nationally representative survey of the middle-aged and older population, has a high-quality data (19).

Conclusion

In conclusion, the frailty index of deficit accumulation is significantly associated with an increased risk of fall among community-dwelling middle-aged and older people in China. It is necessary to integrate the frailty evaluation in routine clinical practice and community-based physical examination of middleaged and older people. Timely screening and recognition of frailty are urgent to prevent falls among middle-aged and older adults.

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