Open Access



Short sleep duration associated with the incidence of cardio-cerebral vascular disease: a prospective cohort study in Shanghai, China

Juzhong Ke¹, Xiaolin Liu¹, Xiaonan Ruan¹, Kang Wu¹, Hua Qiu¹, Xiaonan Wang¹, Zhitao Li¹ and Tao Lin^{1*}

Abstract

Importance Sleep duration plays an important role in predicting CCVD incidence, and have implications for reducing the burden of CCVD. However, the association between sleep duration and predicted cardio-cerebral vascular diseases (CCVD) risk remains to be fully understood.

Objective To investigate the effects of sleep duration on the development of CCVD among Chinese community residents.

Design A prospective cohort study. The baseline survey was conducted from January 2013 to July 2013. The cohort has been followed until December 31, 2016 using a combination of in-person interviews and record linkages with the vital registry of Pudong New Area, Shanghai, China.

Subjects A total of 8245 Chinese community residents were initially enrolled in the cohort. Of those, 6298 underwent the follow-up examination.

Exposure Self-reported sleep duration and sleep quality were obtained via the questionnaire. Sleep duration was divided into five categories: $\leq 5, 6, 7, 8, \text{ or } \geq 9 \text{ h per day.}$

Main Outcome(s) and Measure(s) CCVD, Coronary heart disease (CHD) and Stroke occurrence, Hazard ratios (HRs) and 95% confidence intervals (95% CIs) were calculated using Fine-Gray proportional subdistribution hazards models.

Results During a median follow-up of 3.00 years (IQR 2.92–3.08), we observed 370 participants have had incident CCVD events, of whom 230 had CHDs, 169 had strokes, and 29 had both. After adjustment for relevant confounders, short sleepers (≤ 5 h) had 83% higher risk of total CCVD incidence (HR: 1.83; 95% CI: 1.32–2.54), 82% higher risk of CHD incidence (HR: 1.82; 95% CI: 1.21–2.75), and 82% higher risk of stroke incidence (HR: 1.82; 95% CI: 1.12–2.98) in contrast to the reference group (7 h). Some of these U-shaped relationships varied by age, and were more pronounced in individuals aged < 65 years. Individuals who slept ≤ 5 h per day with baseline hypertension had the highest risk of CCVD incidence (HR: 3.38, 95% CI 2.08–5.48), CHD incidence (HR: 3.11, 95% CI 1.75–5.53), and stroke incidence (HR: 4.33, 95% CI 1.90–9.86), compared with those sleep 7 h and without baseline hypertension.

Conclusions Short sleep duration is independently associated with greater incidence of CCVD, CHD and stroke.

*Correspondence: Tao Lin atolin@yeah.net

Full list of author information is available at the end of the article



© The Author(s) 2023. **Open Access** This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativecommons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

Keywords Sleep duration, Cardio-cerebral vascular disease, Coronary heart disease, Stroke, Prospective cohort studyKey Points

Question What exactly is the relationship between sleep duration and CCVD occurrence in Chinese community residents?

Findings This prospective cohort study investigated the effects of sleep duration on the development of CCVD among 6298 Chinese community residents with a median follow-up of 3.00 years. Short sleepers (≤ 5 h) had higher risk of total CCVD, CHD and stroke incidence compared to the reference group (7 h). Some of these U-shaped relationships varied by age, and were more pronounced in individuals aged<65 years. Individuals who slept ≤ 5 h per day with baseline hypertension had the highest risk of CCVD, CHD and stroke incidence.

Meanings Short sleep duration is independently associated with greater incidence of CCVD, CHD and stroke. Preventative strategies are needed to reduce the disease burden of CCVD associated with inadequate sleep duration.

Introduction

Cardio-cerebral vascular diseases (CCVD) are serious threats to human's health with high incidence, morbidity, mortality, and many complications [1]. Given the severe consequences and accompanying economic losses, recognition of risk factors may play an essential role in preventing CCVD. Extreme sleep duration has long been considered as an important contributing factor to the risk of CCVD [2-4]. Increasing attention has been paid to explore the sleep duration problems and their effects on various health outcomes including CCVD, diabetes mellitus, and cancer [5, 6]. Previous study suggests that between 1985 and 2012, the mean sleep duration has steadily decreased, and the prevalence of short sleep duration (defined as <7 h) has increased from 22.3 to 29.2% [7]. It's critical to understand the role of sleep duration in predicting CCVD incidence, since they may have implications for reducing the burden of CCVD.

Several studies investigated the association between sleep duration and the risk of CCVD outcomes, however, uncertainty still exists about the association and the dose-response relationship [8–11]. In addition, whether the relationship between extreme sleep duration and incident CCVD varies by age or sex remains unclear. If combined effects of sleep duration and hypertension exists on CCVD incidence, increased efforts for their identification and treatment should be undertaken. Although a number of studies have examined such relationships, most of them are cross-sectional designed. In addition, few studies have been done in the Chinese population, where disease burden from CCVD is increasing rapidly in the past decades [12]. Here, we carried out a prospective study to illustrate the association between sleep duration and CCVD incidence among a Chinese population, and to investigate whether this association is age/sex specific and/or modified by hypertension status.

Materials and methods

Study population

This study was performed among subjects of the ongoing prospective population-based Cohort Study in Pudong New Area, Shanghai, China. The study is the sequential research of the previous cross-sectional study [13]. The baseline survey was conducted from January 2013 to July 2013. Of 10,657 subjects in baseline, 1441 were excluded due to the incompletion of the full questionnaire or examination. 971 subjects were excluded because they had been diagnosed CCVD before baseline survey. The remaining 8245 subjects entered the followup cohort. 6220 subjects participated the in-person follow-up survey conducted from January 2016 to May 2016. 78 subjects died before the follow-up surveys, of them 24 died due to CCVD events and 54 due to other causes. 1947 subjects were lost to follow up. Altogether, definite CCVD outcomes of 6298 (76.39%) subjects were obtained, with a median follow-up period of 3.00 years (Fig. 1). The study had obtained approval from the Ethical Committee of Center for Disease Control and Prevention of Pudong New Area. Written informed consents were obtained from all subjects.

Assessment of sleep duration and sleep quality

Self-reported sleep duration and sleep quality were obtained via the questionnaire. Sleep duration was obtained by the question "In the past year, on average, how many hours of sleep do you get in a 24 h period?", answers were divided into five categories: ≤ 5 , 6, 7, 8, or ≥ 9 h. Sleep quality was also inquired. Difficulty in falling asleep was assessed with the question, "In the past year, how often did you have trouble in falling asleep?" Difficulty in staying asleep with the question, "In the past year, how often did you wake up in the midnight and feel difficult to fall asleep again?" Early morning awakening with the question, "In the past year, how often did you wake up in the morning?" Usage of hypnotics with the question, "In the past year, how often did you



Fig. 1 Diagram of this study

use hypnotics or other drugs to help you falling asleep?" Patients with responses of "sixteen to thirty times a month" of at least one of the four items were classified as having poor sleep quality [14].

Assessment of covariates

Information sociodemographic characteristics on (including age, sex, and marital status), smoking, physical activity, alcohol and tea consumption were collected through a questionnaire by face-to-face interview with trained interviewers. Smoking was classified as current smoker (with at least one cigarette a day for at least the previous 6 month) or not. Physical activity was classified as regular exerciser (participating in sports activities for at least once per week in the previous 5 years) or not. Alcohol consumption and tea consumption were classified as regular drinker (at least three times per week for at least the previous 6 month) or not. Standard procedures were used to measure height, weight, SBP and DBP. Body mass index (BMI) was calculated as weight (kg)/ height (m [2]). Hypertension was defined as SBP \geq 140 mmHg or DBP≥90 mmHg, or self-reported history of hypertension. Type 2 diabetes referred to self-reported history of type 2 diabetes, or elevated fast plasma glucose (\geq 7.0mmol/L), or elevated 2-hours plasma glucose (\geq 11.1mmol/L) during 75-g oral glucose tolerance test. Dyslipidemia was defined as LDL-C \geq 4.13mmol/L, or TC \geq 6.20mmol/L, or HDL-C<1.03mmol/L, or TG \geq 2.26mmol/L, or self-reported history of dyslipidemia. The serum C-reactive protein (CRP) levels were detected by particle enhanced immunonephelometric method.

Outcomes

The primary outcome in the current study was CCVD incidence, defined as a physician diagnosis or death of coronary heart disease (CHD) or stroke that occurred in the follow-up duration. Subjects were defined as having CHD if they had angina, myocardial infarction, silent myocardial ischemia or ischemic heart failure, and as having stroke if they had ischemic or hemorrhagic stroke. Deaths were identified through records linking with the vital registry of Pudong New Area, Shanghai, China. Underlying death causes were coded according to ICD-10: I21-I25 for CHD deaths, I60-I64 for stroke deaths. Both CHD and stroke were considered CCVD.

Statistical analysis

Statistical analyses were performed using SPSS 22.0 and SAS 9.4. Subjects were classified into five groups based on sleep duration. Chi-square and one-way ANOVA tests were used to compare the categorical and numeric variables among groups. Taking competing-risks into consideration, Fine-Gray proportional subdistribution hazards model was performed to evaluate the associations of sleep duration with the risk of incident CCVD, covariates were choose based on evidence from literature. Potential confounders included age, sex, marital status, current smoking, alcohol consumption, tea consumption, physical activity, hypertension, type 2 diabetes, dyslipidemia, BMI, CRP, and sleep quality. nonlinear trends of incident CCVD risk were tested by restricted cubic spline Cox regression using 3 knots placed at the 5th, 50th, and 95th percentiles of sleep duration, with sleep duration of 7 h as the reference group [15]. We analyzed the potential interactions by adding interaction terms of age and sex with sleep duration, respectively. We conducted subgroup analyses separately by age group (above and below 65 years). Combined effects of sleep duration and hypertension on the risk of incident CCVD were also evaluated, taking sleep duration of 7 h and non-hypertension as the reference category. The results were presented as hazards ratios (HRs) with 95% confidence intervals (CIs), p < 0.05were considered statistically significant.

Results

Baseline characteristics

Baseline characteristics of subjects are presented in Table 1. Of the 6298 subjects in the study population, 682 subjects (10.83%) reported sleep duration \leq 5 h/day, 1282 (20.36%) slept 6 h/day, 1522 (24.17%) slept 7 h/day,

Table 1 Baseline characteristics stratified by sleep duration

2185 (34.69%) slept 8 h/day, and 627 (9.96%) slept ≥ 9 h/day. Compared to those who slept 7 h a day, those with shorter sleep duration were more likely to be older, female and unmarried. Shorter sleepers were more likely to be diagnosed with hypertension, type 2 diabetes, dyslipidemia and suffered poorer sleep quality; and less likely to be current smoker or tea drinker.

Association between sleep duration and CCVD incidence

During a median follow-up of 3.00 years (IQR 2.92-3.08), we observed 370 participants have had incident CCVD events, of whom 230 had CHDs, 169 had strokes, and 29 had both. 24 individuals died due to fatal CCVD events, including 16 CHDs and 8 strokes. After adjustment for relevant confounders, short sleepers (≤ 5 h) had 83% higher risk of total CCVD incidence (HR: 1.83; 95% CI: 1.32–2.54), 82% higher risk of CHD incidence (HR: 1.82; 95% CI: 1.21-2.75), and 82% higher risk of stroke incidence (HR: 1.82; 95% CI: 1.12-2.98) in contrast to the reference group (7 h). Although both short and long sleepers tended to have a higher risk of CHD incidence, long sleep duration (≥ 9 h) was not associated with total CCVD incidence (HR: 1.20; 95% CI 0.82-1.75), CHD incidence (HR: 1.37; 95% CI 0.86-2.17), or stroke incidence (HR: 1.15; 95% CI 0.65-2.04) (Table 2). The nonlinear associations of sleep duration with risk of incident CCVD were illustrated by the restricted multivariable cubic spline plots as shown in Fig. 2. Multivariable spline analyses showed U-shaped trends in CCVD, CHD and stroke incidence with bottom point at 7 h. Compared to subjects with 7 h sleep duration, shorter sleepers were at higher risk of CCVD, CHD and stroke incidence (*p*<0.001, = 0.002, 0.042, respectively).

	Sleep duration (h)					Total	р
	≤5	6	7	8	≥9	(n=6298)	
	(n=682)	(n=1282)	(n=1522)	(n=2185)	(n=627)	_	
Age (years)	62.01 (9.68)	60.77 (10.50)	57.65 (11.44)	56.26 (12.01)	57.42 (12.06)	58.25 (11.53)	<0.001
Male (%)	204 (29.91%)	435 (33.93%)	583 (38.3%)	879 (40.23%)	233 (37.16%)	2334 (37.06%)	< 0.001
Married (%)	595 (87.24%)	1134 (88.46%)	1383 (90.87%)	1975 (90.39%)	556 (88.68%)	5643 (89.60%)	< 0.001
Current smoking (%)	92 (13.49%)	185 (14.43%)	276 (18.13%)	416 (19.04%)	123 (19.62%)	1092 (17.34%)	< 0.001
Alcohol consumption (%)	80 (11.73%)	137 (10.69%)	185 (12.16%)	296 (13.55%)	80 (12.76%)	778 (12.35%)	0.161
Tea consumption (%)	138 (20.23%)	329 (25.66%)	411 (27.00%)	630 (28.83%)	183 (29.19%)	1691 (26.85%)	< 0.001
Physical activity (%)	189 (27.71%)	367 (28.63%)	400 (26.28%)	557 (25.49%)	165 (26.32%)	1678 (26.64%)	0.331
Hypertension (%)	304 (44.57%)	595 (46.41%)	619 (40.67%)	865 (39.59%)	243 (38.76%)	2626 (41.70%)	< 0.001
Type 2 diabetes (%)	150 (21.99%)	276 (21.53%)	287 (18.86%)	390 (17.85%)	129 (20.57%)	1232 (19.56%)	0.030
Dyslipidemia (%)	365 (53.52%)	617 (48.13%)	666 (43.76%)	1007 (46.09%)	309 (49.28%)	2964 (47.06%)	< 0.001
Poor sleep quality (%)	99 (14.52%)	57 (4.45%)	24 (1.58%)	30 (1.37%)	13 (2.07%)	223 (3.54%)	< 0.001
BMI (kg/m²)	24.92 (3.31)	25.12 (3.44)	25.18 (4.61)	25.04 (3.41)	24.94 (3.66)	25.07 (3.75)	0.487
CRP (mg/L)	0.27 (0.07, 0.93)	0.25 (0.07, 0.90)	0.23 (0.07, 0.87)	0.24 (0.07, 0.94)	0.27 (0.07, 1.01)	0.25 (0.07, 0.94)	0.611

Data are n (%), mean (SD) or median (IQR).

BMI: body mass index, CRP: C-reactive protein

Sleep	Person-years	Inci-	Incidence (per	Hazard	р					
dura-		dent	1000 person-	ratio						
tion		cases	years; 95% CI)	(95% CI)						
(h)										
CCVD										
≤5	1949	73	37.45	1.83	< 0.001					
			(29.02–45.88)	(1.32–2.54)						
6	3748	74	19.74	0.97	0.848					
			(15.29–24.20)	(0.71–1.33)						
7 (ref.)	4470	81	18.12	1.00						
. ,			(14.21-22.03)							
8	6440	103	15.99	0.94	0.657					
			(12.93-19.06)	(0.70-1.25)						
>9	1840	39	21.19	1.20	0.357					
			(14.61-27.77)	(0.82-1.75)						
CHD										
<5	1996	46	23.05	1.82	0.004					
		10	(16.46-29.63)	(1.21-2.75)	0.001					
6	3786	46	1215	0.98	0.910					
-			(8.66–15.64)	(0.65–1.46)						
7 (ref.)	4516	51	11.29	1.00						
()			(8.21–14.38)							
8	6518	59	9.05 (6.75–11.35)	0.84	0.368					
			, , , , , , , , , , , , , , , , , , ,	(0.58–1.22)						
≥9	1866	28	15.01	1.37	0.185					
			(9.49-20.53)	(0.86-2.17)						
Stroke										
<5	2012	34	16.9	1.82	0.017					
			(11.27–22.53)	(1.12-2.98)						
6	3817	32	8 38 (5 49–11 28)	0.91	0 708					
0	5017	52	0.00 (0.10) 11.20)	(0.57–1.47)	0.700					
7 (ref.)	4538	36	7 93 (5 35–10 51)	1 00						
8	6508	50	7.68 (5.56–9.80)	1.04	0.847					
0	0000	50	, .55 (5.56 9.60)	(0.68–1.60)	0.017					
> 9	1871	17	9 09 (4 79-13 30)	1 1 5	0.629					
_ J	10/1	17	J.UJ (T./ J 13.39)	(0.65_2.04)	0.029					

 Table 2
 Association between sleep duration and cardiovascular

 disease incidence in cohort participants

Hazard ratios and 95% CIs are adjusted for age, sex, marriage status, smoking status, drinking status, tea consumption, physical activity, hypertension, diabetes, dyslipidemia, BMI, CRP and sleep quality

Age-specific association between sleep duration and CCVD

incidence

In stratification and interaction analysis, the U-shaped relationship were more prominent among young and middle-aged individuals. In the subgroup of participants aged less than 65 years old, compared with those who slept 7 h per day, those slept \leq 5 h had 2.13 times more risk of incident CHD (HR: 2.13; 95% CI 1.23–3.68), and those slept \geq 9 h had 87% increased risk of incident CHD (HR: 1.87; 95% CI 1.03–3.38), whereas the U-shaped relationship was not found in the subgroup of participants aged more than 65 years old (*p* for interaction=0.032). The results were similar for CCVD incidence, but no significant interaction was found (Fig. 3). The interactions between sleep duration and sex were also analyzed, no significant interaction was observed (all *p* for interaction>0.05).

Combined effects of sleep duration and hypertension on CCVD incidence

In this study, individuals with hypertension had significantly elevated risk of CCVD incidence (HR: 1.67, 95% CI 1.35-2.07), CHD incidence (HR: 1.59, 95% CI 1.21-2.08), and stroke incidence (HR: 1.93, 95% CI 1.40–2.65), relative to non-hypertension individuals. We further analyzed the combined effects of sleep duration and hypertension on CCVD incidence. Individuals who slept ≤ 5 h per day with hypertension had the highest risk of CCVD incidence (HR: 3.38, 95% CI 2.08-5.48), CHD incidence (HR: 3.11, 95% CI 1.75-5.53), and stroke incidence (HR: 4.33, 95% CI 1.90-9.86), compared with those sleep 7 h and without hypertension. Among non-hypertension individuals, elevated risk of CCVD incidence and stroke incidence was observed in ≤ 5 h sleepers, whereas risk of CHD incidence in ≤ 5 h sleepers was elevated but not statistically significant (Fig. 4).



Fig. 2 Multivariable adjusted spline curves for association between sleep duration and risk for (A) CCVD incidence, (B) CHD incidence, and (C) stroke incidence, adjusted for age, sex, marital status, current smoking, alcohol consumption, tea consumption, physical activity, hypertension, type 2 diabetes, dyslipidemia, BMI, CRP, and sleep quality



Fig. 3 Hazard ratios (95% CI) for incident CCVD(A), CHD(B), and Stroke(C) in subjects above and below 65 years according to sleep duration. The adjusted covariates included age, sex, marital status, current smoking, alcohol consumption, tea consumption, physical activity, hypertension, type 2 diabetes, dyslipidemia, BMI, CRP, and sleep quality, with sleep duration of 7 h as the reference group



Fig. 4 Joint association of sleep duration and baseline hypertension status with (A) CCVD incidence, (B) CHD incidence, and (C) stroke incidence. All HRs and 95% Cls were computed with sleep duration of 7 h and absence of hypertension as the reference group, adjusted for age, sex, marital status, current smoking, alcohol consumption, tea consumption, physical activity, hypertension, type 2 diabetes, dyslipidemia, BMI, CRP, and sleep guality

Discussion

In this prospective cohort study, we found that after controlling for a variety of potential confounders, short sleepers (\leq 5 h per day) had higher risk of CCVD, CHD and stroke incidence, and the U-shaped relationship appeared to be seen mainly in individuals less than 65 years old. In addition, there were combined effects of sleep duration and hypertension on incident CCVD risk. This study indicated that maintain a moderate sleep duration may be considered as an effective intervention strategy to prevent CCVD. The study population was randomly recruited from community residents living in Pudong New Area. Pudong New Area has a resident population of about 5 million, accounting for 20% of the total population in Shanghai. In addition, Pudong New Area is the only district with both urban and rural populations in Shanghai, with diverse socioeconomic status, which is a representative model for other populations [16]. Thus, the findings of this study might provide evidence to optimize CCVD control strategy.

This study explored the associations of short and long sleep duration with CCVD incidence based on a Chinese community population, with longitudinal design and multiple potential confounders taken into consideration. In accordance with previous studies, this study adds to the evidence for a U-shaped association of sleep duration with CCVD incidence, and found a robust association between short sleep and increased CCVD risk [2, 6, 17-19]. However, the effects of long sleep duration remain unclear, although some studies reported that sleep duration longer than 8 h/d was associated with an increased risk of CCVD [2, 19, 20], we failed to reach statistical significance in the long sleep group, the result of which was similar to a recent study conducted in Taiwan population [17]. Explanation of the controversial results might be attributed to the diverse races, ages, and lifestyles across studies.

We observed that the increased risk with extreme sleep duration was more prominent in young and middleaged individuals. This age-specific trend is in line with results from a previous study conducted in Australian adults, which showed that short sleep was associated significantly with increased prevalence of heart disease in middle-aged and elderly adults, but not in the very elderly [21]. A study conducted in British population suggested that the increased risk of stroke with short sleep duration was more prominent among middle-aged subjects, but with long sleep more pronounced among the elderly adults [22]. Similarly, the association between sleep duration and hypertension varies by age, Fang J, et al. showed that short sleep was associated with higher likelihood of hypertension among middle-aged adults, but did not among the elderly adults [23]. Although the underlying mechanisms remains unclear, sleep duration might have different implications in different age groups, the increase of age may cover up the influence of sleep duration on CVD incidence among older individuals and leave the detrimental effects robust in younger adults. The interaction by age needs to be tested by larger studies in the future.

Hypertension is the most important risk factor for CCVD, the severe outcome of hypertension is primarily due to its driving role in the development of CCVD [24]. Lower sleeping times are associated with an increased risk of subclinical atherosclerosis, which could be induced by hypertension [25, 26]. Meanwhile, as a chronic disease hallmarked by chronic inflammation, atherosclerosis development has been shown to be potential mechanisms for CCVD development [27]. In this study, the greatest risk of CCVD incidence was found in individuals with both hypertension and short sleep duration, suggesting that appropriate sleep duration is particularly important for patients with hypertension.

As a potentially important confounder, sleep quality has been found to moderate the association between sleep duration and cardiovascular outcomes [28]. Even after excluding subjects with cardiovascular diseases and depression, symptoms of poor sleep quality such as difficulty initiating sleep and non-restorative sleep are associated with a modestly higher risk of CCVD mortality [29]. Therefore, when we investigating the linkage of sleep duration with increased CCVD risk, effects of sleep quality should be ruled out. It is noteworthy that the role of individual differences and their preferences in terms of sleep habits should be taken into consideration [18]. Short sleep duration is often reported in conjunction with lower socioeconomic status and depressive mood, which could bias the link between short sleep and cardiovascular risk [30, 31]. High-risk behaviors such as smoking, drinking, and low physical activity are also prevalent in short sleepers [23]. Adjusting for these established CCVD risk factors did not considerably change the estimated association in this study.

Sleep duration may be involved in the pathogenesis of CCVD, as numerous studies have suggested the relationship between sleep duration and obesity, diabetes as well as increased levels of CRP [32-34]. The mechanisms that underlie these associations are still under investigation. Short sleep duration may lead to sympathetic nervous system overactivity and changes in circadian rhythm, resulting in the rise of blood pressure and arterial stiffness, which may promote cardiovascular risk [28, 35]. Besides, sleep loss impairs glucose homeostasis and insulin sensitivity, leads to health outcomes related to metabolic systems, such as obesity and type 2 diabetes, both plays an important role in development of CCVD [36, 37]. In addition, sleep deprivation has been reported to be associated with elevated levels of inflammatory cytokines, such as CRP and interleukin-6, which is likely to increase CCVD risk through damaging the body's immune function [34]. Proteinuria marks an increased risk of renal damage related to CCVD. Results from observational studies revealed that short sleep duration is associated with prevalence of proteinuria in a dosedependent manner [38, 39]. In patients with chronic kidney disease, short sleep duration is associated with faster decline in renal function and progression to end stage renal disease [40, 41]. Thus, proteinuria might also intermediate the association between short sleep duration and CVD. Further research is needed to clarify these potential roles.

The strengths of this study are its prospective design, relatively large sample size, and the adjustment of many potential risk factors. The findings of this study may provide some new insight into CCVD prediction and prevention. This study has several limitations. First, information about sleep duration were obtained through questionnaires. Sleep duration was a kind of lifestyle factor that might be affected by many factors, thus exposure misclassification bias and discrepancy between measures of subjective sleep duration and objective sleep duration might exist. Previous study suggested that objective sleep duration derived from actigraphy was significantly correlated with subjective self-report sleep duration, but was about 0.5-1 h shorter than the subjective one [42]. Therefore, we may have underestimated the health implications of short sleep duration. Second, the follow-up period of this study was relatively short. Third, although a range of potential confounders were included in the analysis, we could not rule out the possibility of residual/unmeasured confounding.

Conclusions

Short sleep duration is associated with greater incidence of CCVD, independent of other traditional CCVD risk factors. This association appeared to be seen mainly in individuals less than 65 years old. The strongest association with CCVD incidence was found in short sleepers with hypertension. Implementation of preventative

strategies may be needed to reduce the disease burden of CCVD associated with inadequate sleep duration.

Acknowledgements

Not applicable.

Authors' contributions

Drs Ke, and Lin had full access to all of the data in the study, take responsibility for the integrity of the data and the accuracy of the data analysis. Concept and design: Lin

Acquisition, analysis, or interpretation of data: All authors

Drafting of the manuscript: Ke

Critical revision of the manuscript for important intellectual content: All authors

Statistical analysis: Ke

Obtained funding: Ke

Administrative, technical, or material support: Lin Supervision: Lin

Funding

This study was funded by the Young Medical Talents Training Program of the Pudong Health Bureau of Shanghai (PWRq2021-05) to Ke. The funder had no role in the design and conduct of the study; collection, management, analysis, and interpretation of the data; preparation, review, or approval of the manuscript; and decision to submit the manuscript for publication.

Data Availability

All data generated or analyzed during this study are included in this published article.

Declarations

Ethics approval and consent to participate

The study had obtained approval from the Ethical Committee of Center for Disease Control and Prevention of Pudong New Area. Written informed consents were obtained from all subjects. For dead participants, written informed consents were obtained from their legal representatives in addition. The authors declare that all methods were carried out in accordance with relevant guidelines and regulations.

Consent for publication

Not applicable.

Competing interests

The authors declare that they have no competing interests.

Author details

¹Pudong New Area Center for Disease Control and Prevention, Pudong Institute of Preventive Medicine, Fudan University, 3039 Zhangyang Rd, Shanghai 200136, P. R. China

Received: 21 September 2022 / Accepted: 25 March 2023 Published online: 31 March 2023

References

- Thomas H, Diamond J, Vieco A, et al. Global Atlas of Cardiovascular Disease 2000–2016: the path to Prevention and Control. Glob Heart. 2018;13(3):143–63.
- Li W, Wang D, Cao S, et al. Sleep duration and risk of stroke events and stroke mortality: a systematic review and meta-analysis of prospective cohort studies. Int J Cardiol. 2016;223:870–6.
- Yang X, Chen H, Li S, Pan L, Jia C. Association of Sleep Duration with the morbidity and mortality of coronary artery disease: a Meta-analysis of prospective studies. Heart Lung Circ. 2015;24(12):1180–90.
- Cappuccio FP, Cooper D, D'Elia L, Strazzullo P, Miller MA. Sleep duration predicts cardiovascular outcomes: a systematic review and meta-analysis of prospective studies. Eur Heart J. 2011;32(12):1484–92.

- Cai H, Shu XO, Xiang YB, et al. Sleep duration and mortality: a prospective study of 113 138 middle-aged and elderly chinese men and women. Sleep. 2015;38(4):529–36.
- Yin J, Jin X, Shan Z et al. Relationship of Sleep Duration With All-Cause Mortality and Cardiovascular Events: A Systematic Review and Dose-Response Meta-Analysis of Prospective Cohort Studies. J Am Heart Assoc. 2017;6(9).
- 7. Ford ES, Cunningham TJ, Croft JB. Trends in Self-Reported sleep duration among US adults from 1985 to 2012. Sleep. 2015;38(5):829–32.
- Pan A, De Silva DA, Yuan JM, Koh WP. Sleep duration and risk of stroke mortality among chinese adults: Singapore Chinese health study. Stroke. 2014;45(6):1620–5.
- Kim Y, Wilkens LR, Schembre SM, Henderson BE, Kolonel LN, Goodman MT. Insufficient and excessive amounts of sleep increase the risk of premature death from cardiovascular and other diseases: the multiethnic cohort study. Prev Med. 2013;57(4):377–85.
- Xiao Q, Keadle SK, Hollenbeck AR, Matthews CE. Sleep duration and total and cause-specific mortality in a large US cohort: interrelationships with physical activity, sedentary behavior, and body mass index. Am J Epidemiol. 2014;180(10):997–1006.
- Hausler N, Lisan Q, Van Sloten T et al. Cardiovascular health and sleep disturbances in two population-based cohort studies. Heart. 2019 Oct;105(19):1500–1506.
- Liu S, Li Y, Zeng X et al. Burden of Cardiovascular Diseases in China, 1990– 2016: Findings From the 2016 Global Burden of Disease Study. Jama Cardiol. 2019 Apr1;4(4):342–352.
- Ke J, Zhou X, Qiu H, et al. Sex-specific associations between extreme sleep duration and prevalence of cardio-cerebral vascular disease: a communitybased cross-sectional study. Sleep Med. 2018;42:61–7.
- Bertisch SM, Pollock BD, Mittleman MA et al. Insomnia with objective short sleep duration and risk of incident cardiovascular disease and all-cause mortality: Sleep Heart Health Study. Sleep. 2018;41(6).
- Cappuccio FP, Cooper D, D'Elia L, Strazzullo P, Miller MA. Sleep duration predicts cardiovascular outcomes: a systematic review and meta-analysis of prospective studies. Eur Heart J. 2011;32(12):1484–92.
- Ke J, Lin T, Liu X, et al. Glucose intolerance and Cancer Risk: A Community-Based prospective cohort study in Shanghai, China. Front Oncol. 2021;11:726672.
- Lao XQ, Liu X, Deng HB, et al. Sleep Quality, Sleep Duration, and the risk of Coronary Heart Disease: a prospective cohort study with 60,586 adults. J Clin Sleep Med. 2018;14(1):109–17.
- Itani O, Jike M, Watanabe N, Kaneita Y. Short sleep duration and health outcomes: a systematic review, meta-analysis, and meta-regression. Sleep Med. 2017;32:246–56.
- Wang D, Li W, Cui X, et al. Sleep duration and risk of coronary heart disease: a systematic review and meta-analysis of prospective cohort studies. Int J Cardiol. 2016;219:231–9.
- Jike M, Itani O, Watanabe N, Buysse DJ, Kaneita Y. Long sleep duration and health outcomes: a systematic review, meta-analysis and meta-regression. Sleep Med Rev. 2018;39:25–36.
- Magee CA, Kritharides L, Attia J, McElduff P, Banks E. Short and long sleep duration are associated with prevalent cardiovascular disease in australian adults. J Sleep Res. 2012;21(4):441–7.
- 22. Leng Y, Cappuccio FP, Wainwright NW, et al. Sleep duration and risk of fatal and nonfatal stroke: a prospective study and meta-analysis. Neurology. 2015;84(11):1072–9.
- 23. Fang J, Wheaton AG, Keenan NL, Greenlund KJ, Perry GS, Croft JB. Association of sleep duration and hypertension among US adults varies by age and sex. Am J Hypertens. 2012;25(3):335–41.
- 24. Joyner MJ, Limberg JK. Blood pressure: return of the Sympathetics? Curr Hypertens Rep. 2016;18(1):7.
- Dominguez F, Fuster V, Fernandez-Alvira JM, et al. Association of Sleep Duration and Quality with subclinical atherosclerosis. J Am Coll Cardiol. 2019;73(2):134–44.
- Blasco-Colmenares E, Moreno-Franco B, Latre ML, et al. Sleep duration and subclinical atherosclerosis: the Aragon Workers' Health Study. Atherosclerosis. 2018;274:35–40.
- 27. Pothineni N, Subramany S, Kuriakose K, et al. Infections, atherosclerosis, and coronary heart disease. Eur Heart J. 2017;38(43):3195–201.
- Covassin N, Singh P. Sleep Duration and Cardiovascular Disease Risk: epidemiologic and experimental evidence. Sleep Med Clin. 2016;11(1):81–9.

- Wu MP, Lin HJ, Weng SF, Ho CH, Wang JJ, Hsu YW. Insomnia subtypes and the subsequent risks of stroke: report from a nationally representative cohort. Stroke. 2014;45(5):1349–54.
- Sands-Lincoln M, Loucks EB, Lu B, et al. Sleep duration, insomnia, and coronary heart disease among postmenopausal women in the Women's Health Initiative. J Womens Health (Larchmt). 2013;22(6):477–86.
- 32. Wu Y, Zhai L, Zhang D. Sleep duration and obesity among adults: a metaanalysis of prospective studies. Sleep Med. 2014;15(12):1456–62.
- Shan Z, Ma H, Xie M, et al. Sleep duration and risk of type 2 diabetes: a metaanalysis of prospective studies. Diabetes Care. 2015;38(3):529–37.
- Chiang JK. Short duration of sleep is associated with elevated high-sensitivity C-reactive protein level in taiwanese adults: a cross-sectional study. J Clin Sleep Med. 2014;10(7):743–9.
- Sunbul M, Kanar BG, Durmus E, Kivrak T, Sari I. Acute sleep deprivation is associated with increased arterial stiffness in healthy young adults. Sleep Breath. 2014;18(1):215–20.
- Deng HB, Tam T, Zee BC et al. Short Sleep Duration Increases Metabolic Impact in Healthy Adults: A Population-Based Cohort Study. Sleep. 2017;40(10).

- 37. Maskarinec G, Jacobs S, Amshoff Y, et al. Sleep duration and incidence of type 2 diabetes: the multiethnic cohort. Sleep Health. 2018;4(1):27–32.
- Aoki K, Yamamoto R, Shinzawa M, et al. Sleep debt and prevalence of proteinuria in subjects with short sleep duration on weekdays: a cross-sectional study. Clin Exp Nephrol. 2020;24(2):143–50.
- Cheungpasitporn W, Thongprayoon C, Gonzalez-Suarez ML, et al. The effects of short sleep duration on proteinuria and chronic kidney disease: a systematic review and meta-analysis. Nephrol Dial Transplant. 2017;32(6):991–6.
- Yamamoto R, Shinzawa M, Isaka Y, et al. Sleep Quality and Sleep Duration with CKD are Associated with Progression to ESKD. Clin J Am Soc Nephrol. 2018;13(12):1825–32.
- 41. McMullan CJ, Curhan GC, Forman JP. Association of short sleep duration and rapid decline in renal function. Kidney Int. 2016;89(6):1324–30.
- 42. Matsumoto T, Chin K. Prevalence of sleep disturbances: sleep disordered breathing, short sleep duration, and non-restorative sleep. Respir Investig. 2019;57(3):227–37.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.