# Association of accelerometer-measured physical activity with kidney function in a Japanese population: the DOSANCO Health Study <br> Check for updates 

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#### Abstract

Background: Sedentary behavior and decreased physical activity are associated with reduced kidney function, yet most evidence is based on self-reported physical activity. This study investigated the association between accelerom-eter-based physical activity level and kidney function in a general Japanese population. Methods: A cross-sectional study was conducted in 440 community-dwelling Japanese participants, aged 35-79years. Time ( $\mathrm{min} / \mathrm{d}$ ) was assessed for the following types of physical activity: sedentary behavior, light physical activity (LPA), and moderate-to-vigorous physical activity (MVPA). Kidney function was assessed using estimated glomerular filtration rate (eGFR). A linear regression model was employed to calculate the $\beta$ coefficient of eGFR for a $60-\mathrm{min} / \mathrm{d}$ increase in sedentary behavior and LPA and a $10-\mathrm{min} / \mathrm{d}$ increase in MVPA. A logistic regression model was used to calculate the odds ratio for low eGFR ( $<60$ versus $\geq 60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ) for a $60-\mathrm{min} / \mathrm{d}$ or $10-\mathrm{min} / \mathrm{d}$ increase in each physical activity type. Results: MVPA time and eGFR were positively associated in both men and women, after adjusting for age, body mass index, and other clinical characteristics (Men: $\beta, 0.91 ; P=0.021$; Women: $\beta, 0.70 ; P=0.034$ ). In women, sedentary behavior and eGFR were inversely associated after adjusting for the same factors ( $\beta,-1.06 ; P=0.048$ ). The odds ratio ( $95 \%$ confidence interval) for low eGFR associated with a 60 -min increase in sedentary behavior was 1.65 (1.07-2.55) after adjusting for the same factors in women.

Conclusion: Longer sedentary behavior and shorter MVPA time were associated with lower kidney function in the Japanese population.


Keywords: Estimated glomerular filtration rate, Physical activity, Sedentary behavior, Accelerometer
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## Introduction

Chronic kidney disease (CKD) is a public health concern worldwide [1, 2]. Previous studies have found that reduced kidney function, even within the range of nonCKD, is a major risk factor for cardiovascular events and deaths independent of classical risk factors, including diabetes and hypertension [3-5]. To reduce this excess burden of CKD, the identification and intervention of
modifiable risk factors for the initiation and progression of CKD at an earlier disease stage are important [6].
Decreased physical activity is a modifiable risk factor for cardiovascular disease [7]. Recently, an interest has developed in sedentary behavior that differs from a lack of physical activity, with respect to the risk of cardiovascular disease [8, 9], as well as CKD. Several cross-sectional studies have reported that self-reported sedentary behavior, often independent of reduced time spent engaging in moderate-to-vigorous physical activity (MVPA), was associated with reduced kidney function [10-12]. Although questionnaires assess a variety of physical activity patterns, the use of questionnaires in general is unreliable in assessing the amount of time spent in light physical activity (LPA) and sedentary behavior, leading to the misclassification of inactivity [13, 14]. In this regard, an accelerometer provides a detailed assessment of the quantity and quality of physical activity. However, very few studies have examined the association between accelerometer-based physical activity level and kidney function in the general population of Western countries [14, 15]. This relationship remains unclear in general Asian populations that have different cardiovascular risk profiles compared to the Western populations [16]. Therefore, the present study investigated the association between indices of physical activity measured with an accelerometer and kidney function in a general Japanese population. The accelerometer measured both physical activity intensity and time.

## Methods

## Study design and population

We conducted a cross-sectional study as part of the Dynamics of Lifestyle and Neighborhood Community on Health Study (DOSANCO Health Study), a communitybased study in Suttu, Hokkaido prefecture, Japan, from May 2015 to November 2015 [17]. This study invited all residents aged 3 years or above (as criterion for inclusion) in Suttu, except for those living in nursing homes (as criterion for exclusion). Briefly, a total of 2638 residents aged $3-103$ years and living at home were considered eligible participants for inclusion, and 2100 participants ( 977 men and 1123 women) completed a self-administered questionnaire. Of 1379 ( 650 men and 729 women) participants ages 35-79 for whom we undertook an additional detailed survey, 545 participants ( 245 men and 300 women) also provided a blood sample. At the same time, these participants were further asked to wear an accelerometer for 14 days.
For the present study, we restricted our analyses to 488 participants who provided valid data on an accelerometer (i.e., 4 days of at least 10 h of consecutive wearing time) [18]. Of the 488 participants, 48 were excluded
for the following reasons: history of myocardial infarction ( $n=16$ ) or stroke ( $n=17$ ), chronic obstructive pulmonary disease ( $n=2$ ), and missing data on variables other than accelerometer indices and serum cystatin C level $(n=13)$. Therefore, a total of 440 participants (190 men and 250 women) were included in the final analysis (Fig. 1).

## Assessment of physical activity

An accelerometer (Active Style Pro HJA-750C; Omron Healthcare, Kyoto, Japan) was used to assess physical activity for 14 days. This devise is validated [19, 20], and comparable to other devices commonly used in epidemiological studies [21, 22]. In short, the Active Style Pro has the dimensions of $40 \times 52 \times 12 \mathrm{~mm}$, weighs 23 g , and records anteroposterior ( $x$-axis), mediolateral ( $y$-axis), and vertical (z-axis) accelerations with a resolution of 3 mG at 32 Hz . Before assessment, the accelerometer was calibrated according to the manufacturer's recommendations. Each participant wore an accelerometer on the left hip for 14 days while awake, and removed it for water activities and bathing. Non-wear time was defined as 60 mins of continuous zero counts [23], and a standard 60 -s data period was used [24, 25]. Valid accelerometer data was obtained if the device was worn for $\geq 10 \mathrm{~h} / \mathrm{d}$ for at least 4 days [26]. The accelerometer provided data on the intensity of physical activity expressed as metabolic equivalents (METs) using equations for estimation [20, 27]. Time ( $\mathrm{min} / \mathrm{d}$ ) was assessed for each of the following three physical activity intensities based on METs: sedentary behavior, $\leq 1.5 \mathrm{METs}$; LPA, 1.6-2.9 METs; and MVPA, $\geq 3.0$ METs [20, 28].

## Assessment of kidney function

Blood samples were collected after overnight fast from the antecubital vein of seated participants with minimal tourniquet use. Serum was separated and centrifuged after blood coagulation, and stored at $-80^{\circ} \mathrm{C}$ until cystatin $C$ level was measured. Serum cystatin $C$ level was measured on a latex immunoturbidimetric assay using an automated analyzer (JCA-BM8060; JEOL, Tokyo, Japan) in a single laboratory. Estimated glomerular filtration rate (eGFR) was calculated using the Japan Society of Nephrology Chronic Kidney Disease Initiative (JSN-CKDI) equation with serum cystatin C [29]. Serum cystatin C level has an advantage of being less influenced by various factors including age, sex, muscle mass, diet, and physical activity, in contrast to serum creatinine [30].

## Assessment of other factors

Blood pressure was measured twice using an automatic manometer (USM-700G Si; Ueda Avancer Corp., Tokyo, Japan), after the participant rested for 5 min in


Fig. 1 Study flow chart for inclusion of the participants
the seated position. The mean of the first and second readings was used for the analysis. Body height and weight were measured, and body mass index (BMI) was calculated as weight ( kg )/height $\left(\mathrm{m}^{2}\right)$. Enzymatic assays were used to measure levels of total cholesterol and high-density lipoprotein (HDL) cholesterol. NonHDL cholesterol was calculated as total cholesterol minus HDL cholesterol. Hemoglobin A1c (HbA1c) was measured using a latex agglutination immunoassay (Hitachi Chemical Diagnostics Systems Co., Ltd., Tokyo, Japan). Hypertension was defined as either systolic blood pressure $\geq 140 \mathrm{mmHg}$, diastolic blood pressure $\geq 90 \mathrm{mmHg}$, or use of antihypertensive medication [31]. Diabetes was defined as either fasting blood glucose level $\geq 126 \mathrm{mg} / \mathrm{dL}, \mathrm{HbA1c}>6.5 \%$, or self-reported use of diabetes medication [32]. Dyslipidemia was defined as either non-HDL-C $\geq 170 \mathrm{mg} / \mathrm{dL}$ or use of medication for dyslipidemia [33]. A self-administered questionnaire collected data, including age, sex, medical history, smoking status, alcohol drinking, dietary intake, and educational background. Smoking status was categorized as "never smoker," "ex-smoker," or "current smoker." Habitual alcohol drinking was categorized as "non-drinker," "ex-drinker," or "current drinker." Dietary intake was assessed using a brief self-administered diet history questionnaire [34]. The data were subsequently converted into total energy intake ( $\mathrm{kcal} / \mathrm{d}$ ), fat intake ( $\mathrm{g} / \mathrm{d}$ ), and sodium intake ( $\mathrm{g} / \mathrm{d}$ ). The education
level was categorized as either "high school or less" or "more than high school."

## Statistical analysis

Data were analyzed separately for men and women due to sex differences in sedentary behavior and physical activity [25, 35]. Initially, we examined the association between time of each physical activity type (i.e., sedentary behavior, LPA, and MVPA; min/d as continuous variables) and eGFR ( $\mathrm{mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ as a continuous variable) among all study participants, using a linear regression model. We calculated the $\beta$ coefficient and $95 \%$ confidence interval (CI) of eGFR for i) a $60-\mathrm{min} / \mathrm{d}$ increase in sedentary behavior, ii) a $60-\mathrm{min} / \mathrm{d}$ increase in LPA, and iii) a $10-\mathrm{min} / \mathrm{d}$ increase in MVPA $[36,37]$ using the following models. Model 1 was adjusted for age (years as a continuous variable), accelerometer wear time ( $\mathrm{min} / \mathrm{d}$ as a continuous variable). In contrast, model 2 was adjusted for model 1 variables plus BMI ( $\mathrm{kg} / \mathrm{m}^{2}$ as a continuous variable), smoking status (never, formerly, or currently smoking, using 2 dummy variables with "never smoking" as the reference), alcohol drinking (never, formerly, or currently drinking, using 2 dummy variables with "never drinking" as the reference), energy intake ( $\mathrm{kcal} / \mathrm{d}$ as a continuous variable), fat intake ( $\mathrm{g} / \mathrm{d}$ as a continuous variable), sodium intake ( $\mathrm{g} / \mathrm{d}$ as a continuous variable), educational background ("high school or less" or "more than high
school"), hypertension (no or yes), diabetes (no or yes), and dyslipidemia (no or yes). Model 3 further incorporated time of the other physical activity types to assess whether the association of interest was independent of the other physical activity types [36]. Model 3 was adjusted for model 2 plus MVPA time when sedentary behavior and LPA time were examined, whereas sedentary behavior was incorporated as a covariate when MVPA time was examined. Next, we calculated the odds ratio and $95 \%$ CI of low eGFR ( $<60$ versus $\geq 60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ) [38], for a $60-\mathrm{min} / \mathrm{d}$ increase in sedentary behavior or LPA, and for a $10-\mathrm{min} / \mathrm{d}$ increase in MVPA among all study participants; a logistic regression model that incorporated the same variables as covariates was used. All analyses were conducted using JMP Pro software version 14.0.0 for Macintosh (SAS Institute, Cary, NC, USA). Statistical significance was defined as a two-tailed $P<0.05$.

## Results

Table 1 shows the characteristics of the 440 study participants together and by sex. In the overall sample, the mean age $\pm$ standard deviation was $58.3 \pm 12.1 \mathrm{y}$. The mean length of accelerometer wear was 12 d and wear time was $883.1 \pm 90.0 \mathrm{~min} / \mathrm{d}$. Mean $\pm$ standard deviation of GFR estimated by the JSN-CKDI equation with cystatin C level was $92.6 \pm 19.9 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$, and $5.5 \%$ of all participants ( $5.8 \%$ men and $5.2 \%$ women) had low eGFR $\left(<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}\right)$. No participants had eGFR $<30 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$. Women had lower mean BMI, total energy intake, and sodium intake than men. Women were less likely to be smokers and drinkers, had less education, and were more likely to have dyslipidemia than men. In addition, women had a shorter mean time spent engaging in sedentary behavior and a longer mean of LPA and MVPA time than men.
Figure 2 shows a sex-specific scatter graph of sedentary behavior, LPA, MVPA, and eGFR. Time in MVPA showed

Table 1 Demographic and clinical characteristics of the 440 participants from Suttu town, Hokkaido, Japan

|  | Overall $(n=440)$ | Men $(n=190)$ | Women $(n=250)$ | $p$-value for difference |
| :---: | :---: | :---: | :---: | :---: |
| Age (years) | $58.3 \pm 12.1$ | $57.3 \pm 12.1$ | $59.0 \pm 12.0$ | 0.13 |
| Body mass index ( $\mathrm{kg} / \mathrm{m}^{2}$ ) | $23.7 \pm 3.7$ | $24.4 \pm 3.5$ | $23.2 \pm 3.8$ | < 0.001 |
| Smoking status (\%) |  |  |  | < 0.001 |
| Never smoker | 47.1\% (208) | 17.4\% (33) | 70.0\% (175) |  |
| Former smoker | 32.0\% (141) | 50.0\% (95) | 18.4\% (46) |  |
| Current smoker | 20.7\% (91) | 32.6\% (62) | 11.6\% (29) |  |
| Alcohol drinking (\%) |  |  |  | $<0.001$ |
| Never drinker | 33.9\% (150) | 17.9\% (34) | 46.4\% (116) |  |
| Former drinker | 8.4\% (37) | 10.0\% (19) | 7.2\% (18) |  |
| Current drinker | 57.5\% (253) | 71.1\% (137) | 46.4\% (116) |  |
| Total energy intake (kcal/d) | $1760.7 \pm 539.7$ | $1953.0 \pm 546.2$ | $1614.7 \pm 487.3$ | $<0.001$ |
| Sodium intake (g/d) | $10.9 \pm 3.6$ | $11.7 \pm 3.8$ | $10.3 \pm 3.4$ | $<0.001$ |
| Fat intake (g/d) | $49.8 \pm 19.1$ | $50.9 \pm 18.8$ | $49.0 \pm 19.4$ | 0.31 |
| Educational background |  |  |  | 0.016 |
| < High school | 57.0\% (251) | 50.5\% (96) | 62.0\% (155) |  |
| $\geq$ High school | 43.0\% (189) | 49.5\% (94) | 38.0\% (95) |  |
| Hypertension | 57.7\% (254) | 57.4\% (109) | 58.0\% (145) | 0.89 |
| Diabetes | 14.8\% (65) | 15.8\% (30) | 14.0\% (35) | 0.60 |
| Dyslipidemia | 34.5\% (152) | 27.9\% (53) | 39.6\% (99) | 0.011 |
| eGFR cystatin | $92.6 \pm 19.9$ | $92.4 \pm 19.3$ | $92.8 \pm 20.4$ | 0.82 |
| eGFR< $60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ (\%) | 5.5\% (24) | 5.8\% (11) | 5.2\% (13) | 0.79 |
| Sedentary behavior (min/d) | $463.2 \pm 117.5$ | $494.6 \pm 119.9$ | $439.4 \pm 110.0$ | $<0.001$ |
| Light physical activity (min/d) | $369.1 \pm 95.6$ | $325.0 \pm 83.5$ | $402.6 \pm 90.6$ | < 0.001 |
| Moderate-to-vigorous physical activity (min/d) | $50.5 \pm 31.1$ | $47.2 \pm 29.9$ | $53.1 \pm 31.8$ | 0.049 |

Data are presented for the total study group and also grouped according to sex
Variables are presented as mean $\pm$ standard deviation, median (interquartile range), or the \% (number) of participants in that category. eGFR estimated glomerular filtration rate. Student's t-test, chi-squared test was used to compare participant characteristics between sex


Fig. 2 Correlations between sedentary behavior/physical activity and estimated glomerular filtration rate (eGFR) in the study population

Table 2 Linear regression coefficient ( $95 \% \mathrm{Cl}$ ) of eGFR for a $60-\mathrm{min} / \mathrm{d}$ increase for sedentary behavior, light physical activity, and for a 10-min/d increase for moderate-to-vigorous physical activity in the 440 participants


Linear regression model was used to calculate coefficient: Model 1 was adjusted for age and accelerometer wear time. Model 2 was adjusted for the same covariates used in model 1 plus body mass index, smoking status, alcohol drinking, total energy intake, sodium intake, fat intake, educational background, hypertension, diabetes, and dyslipidemia. Model 3 was adjusted for the same covariates used in model 2 plus moderate-to-vigorous physical activity (when sedentary behaviour and light activity were examined) or sedentary (when moderate-to-vigorous physical activity was examined)
Abbreviations: eGFR estimated glomerular filtration rate; CI confidence interval
a positive correlation with eGFR in both men and women (men, $\mathrm{r}=0.29, p<0.001$; women, $\mathrm{r}=0.19, p<0.01$ ).
Table 2 shows the $\beta$ coefficients of eGFR for a $60-\mathrm{min} / \mathrm{d}$ increase in sedentary behavior and LPA, and for a $10-\mathrm{min} / \mathrm{d}$ increase in MVPA in men and women. For both men and women, MVPA time was positively associated with eGFR (Men: $\beta, 0.91 ; 95 \%$ CI: $0.14-1.69$; Women: $\beta, 0.70 ; 95 \% \mathrm{CI}: 0.055-1.34$ ) after adjusting for possible confounding factors, including age, accelerometer wear time, BMI, smoking status, alcohol drinking, total energy intake, fat intake, sodium intake, educational background, hypertension, diabetes, and dyslipidemia (model 2). This positive association was observed even after further adjusting for sedentary behavior only in men ( $\beta, 1.02$; 95\% CI: 0.15-1.90) (model 3). In women, sedentary behavior was inversely associated with eGFR after adjusting for the same potential confounders ( $\beta,-1.06$; $95 \% \mathrm{CI} ;-2.11--0.01$ ) (model 2). This inverse association was attenuated after further adjusting for MVPA time ( $\beta,-0.69$; 95\% CI: $-1.86-0.47$ ) (model 3).
Table 3 shows the odds ratios of low eGFR ( $<60 \mathrm{~mL} /$ $\mathrm{min} / 1.73 \mathrm{~m}^{2}$ ) for a $60-\mathrm{min} / \mathrm{d}$ increase in sedentary behavior and LPA and for a $10-\mathrm{min} / \mathrm{d}$ increase in MVPA among men and women. In women, the odds ratio of low eGFR ( $<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ) associated with a $60-\mathrm{min} / \mathrm{d}$
increase in sedentary behavior was 1.65 (95\% CI: 1.072.55 ), after adjusting for possible confounding factors, including age, accelerometer wear time, BMI, smoking status, alcohol drinking, total energy intake, fat intake, sodium intake, educational background, hypertension, diabetes, and dyslipidemia (model 2). The odds ratio was slightly attenuated at 1.51 ( $95 \% \mathrm{CI}: 0.94-2.41$ ), even after further adjusting for MVPA time (model 3).

## Discussion

The present cross-sectional study measured the time spent engaging in sedentary behavior, LPA, and MVPA with an accelerometer among adult residents of a rural community in Japan. Our study showed that time spent engaging in MVPA was positively associated with kidney function (quantified by eGFR with cystatin C) in both men and women, while sedentary behavior time was inversely associated with kidney function in women only. These associations were observed even after accounting for relevant clinical characteristics, including BMI, hypertension, diabetes, and glucose metabolism. Therefore, our study highlighted that the type of physical activity had different associations with reduced kidney function, within clinically normal ranges, for men and women.

Table 3 Odds ratio of low eGFR ( $<60$ versus $\geq 60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ) for a $60-\mathrm{min} / \mathrm{d}$ increase for sedentary behavior, light physical activity, and for a $10-\mathrm{min} / \mathrm{d}$ increase for moderate-to-vigorous physical activity in the 440 participants

|  | Crude |  | Adjusted, model 1 |  | Adjusted, model 2 |  | Adjusted, model 3 |  |
| :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: | :---: |
|  | OR (95\% CI) | $p$-value | OR (95\% CI) | $p$-value | OR (95\% CI) | $p$-value | OR (95\% CI) | $p$-value |
| Men |  |  |  |  |  |  |  |  |
| Sedentary behavior (min/d) | 1.11 (0.82, 1.50) | 0.51 | 1.36 (0.85, 2.17) | 0.21 | 1.58 (0.88, 2.82) | 0.10 | 1.56 (0.78, 3.13) | 0.19 |
| Light physical activity (min/d) | 0.81 (0.50, 1.30) | 0.37 | 0.88 (0.46, 1.70) | 0.71 | 0.89 (0.41, 1.93) | 0.77 | 1.07 (0.47, 2.44) | 0.88 |
| Moderate-to-vigorous physical activity ( $\mathrm{min} / \mathrm{d}$ ) | $0.61(0.40,0.90)$ | 0.023 | 0.84 (0.58, 1.20) | 0.30 | 0.75 (0.48, 1.16) | 0.15 | 0.88 (0.56, 1.37) | 0.55 |
| Women |  |  |  |  |  |  |  |  |
| Sedentary behavior (min/d) | 1.45 (1.09, 1.92) | 0.011 | 1.41 (1.05, 1.94) | 0.021 | 1.65 (1.07, 2.55) | 0.013 | 1.51 (0.94, 2.41) | 0.072 |
| Light physical activity (min/d) | 0.72 (0.49, 1.07) | 0.10 | 0.60 (0.35, 0.97) | 0.037 | 0.60 (0.31, 1.15) | 0.11 | 0.67 (0.33, 1.36) | 0.26 |
| Moderate-to-vigorous physical activity ( $\mathrm{min} / \mathrm{d}$ ) | 0.72 (0.53, 0.93) | 0.0085 | 0.76 (0.54, 1.00) | 0.052 | 0.69 (0.46, 1.04) | 0.036 | 0.77 (0.49, 1.21) | 0.22 |

Logistic regression model was used to calculate odds ratio: Model 1 was adjusted for age and accelerometer wear time. Model 2 was adjusted for the same covariates used in model 1 plus body mass index, smoking status, alcohol drinking, total energy intake, sodium intake, fat intake, educational background, hypertension, diabetes, and dyslipidemia. Model 3 was adjusted for the same covariates used in model 2 plus moderate-to-vigorous physical activity (when sedentary behavior and light activity were examined) or sedentary (when moderate-to-vigorous physical activity was examined). Abbreviations: OR odds ratio; Cl confidence interval

On average, people engage in sedentary behaviors for approximately $55 \%$ of their awake hours [15]. Many epidemiological studies suggest that sedentary behavior increases the risks of cardiovascular disease and all-cause mortality, independent of obesity [8]. However, limited epidemiological evidence is available on the link between sedentary behavior and reduced kidney function, which is a worldwide public health concern [1, 2]. A previous cross-sectional study involving community-dwelling Japanese older adults showed that individuals who engage in sedentary behavior for longer periods of time (as self-reported) had a higher prevalence of low eGFR of $<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ [12]. In a United States cohort study, middle-aged participants who participated in MVPA for $>300 \mathrm{~min} /$ week (as self-reported) had an $11 \%$ reduced risk of incident reduced eGFR of $<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ over a $24-y$ follow-up period, compared with those without the corresponding MVPA time [39]. To date, very few studies have examined the association between a reliable measure of sedentary behavior and physical activity and kidney function. The National Health and Nutrition Examination Survey in the U.S. reported that sedentary behavior measured with an accelerometer was inversely associated with eGFR among non-diabetic women [10]. Although men and women were combined in the analyses of these previous studies, the results are partially in accordance with our results for women, but not for men. Meanwhile, a recent cross-sectional study among older adults in the United Kingdom revealed that every 10-min increase in MVPA time had an odds ratio of 0.87 (95\% CI: 0.78-0.96) for moderately reduced eGFR of $<45 \mathrm{~mL} /$ $\min / 1.73 \mathrm{~m}^{2}$ [36]. The results of this previous study are partially consistent with our results.

Evidence suggests that physical activity has different beneficial effects on cardiovascular disease in men and women [40, 41]. Similar sex differences may be applicable to renal dysfunction by the following biological differences between men and women. A recent report showed a more positive association between time spent participating in sedentary behavior and pro-inflammatory biomarker levels (such as IL-6 and fibrinogen) in women compared to men [42]. Thus, this report may be similar to our results that sedentary behavior is likely to be associated with reduced kidney function in women only.

Patients with CKD and end-stage renal disease are less physically active and have more sedentary behaviors than healthy individuals [43, 44]. A recent nationwide cross-sectional study reported that, even those with slightly reduced eGFR were more likely to have sedentary behavior than those with normal kidney function (adjusted odds ratio 1.7; 95\% CI: 1.2-2.3) [15]. In this study of participants with general population, lower eGFR was associated with decreased MVPA time and increased sedentary behavior. The possible reason for the association between reduced eGFR and less MVPA or increased sitting time is unclear. The Third National Health and Nutrition Examination Survey reported that the incidence of frailty was higher among those with even participants with stage 1 and 2 CKD compared to those without CKD [45]. In addition, a recent cross-sectional study with general population showed that eGFR and skeletal muscle mass per body weight were significantly correlated [46]. It is possible that the physiological changes in the muscles associated with renal dysfunction may decrease physical activity. Meanwhile, according to the 2012 Kidney Disease Improving Global Outcomes
(KDIGO) clinical practice guidelines, exercise is recommended for CKD patients [47]. However, there is limited evidence for the benefits of increased physical activity and decreased sedentary behavior for renal function among patients with early-stage CKD, the best candidates who may benefit from healthy behavioral modifications. A recent cohort study in Korea involving 83,470 subjects with early-stage CKD showed that individuals engaged in sufficient physical activity (as self-reported) had significantly reduced risk of death (adjusted hazard ratio $0.73 ; 95 \%$ CI: $0.64-0.83 ; p<0.001$ ) [48]. The results of our study, which objectively assessed physical activity, are consistent with those of previous studies on patients with early-stage CKD. Although the exact mechanism remains uncertain, longer MVPA time may improve endothelial function of vessels and prevent atherosclerotic change, which may consequently preserve kidney function [49]. Further research is needed to clarify the benefits of increased physical activity for renal function among healthy individuals with normal renal function.
Although our analysis showed that LPA time was not related to renal function, a previous epidemiological study had contrasting results. The study examined the relationship between subjectively measured physical activity and the odds of low eGFR ( $<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ), and it suggested that increased LPA time was associated with lower odds of low eGFR [37]. One possible reason for the varied findings between our study and the previous study is the difference in the MVPA time. In the previous studies, the protective effects of increased LPA on mortality were limited to people with low MVPA times [40, 50]. Our study participants had a mean MVPA time of $47.0 \mathrm{~min} / \mathrm{d}$ for men and $53.1 \mathrm{~min} / \mathrm{d}$ for women. Thus, we may have observed a null association between LPA time and kidney function for both men and women. It may not be surprising that our results regarding LPA contradict those of a previous study, which showed a positive association between LPA time and kidney function among older adults; the participants had a lower mean MVPA time compared to our study participants [36].
The strength of our study is that we objectively measured sedentary behavior and physical activity using an accelerometer, which allowed us to accurately estimate the time spent in various intensities of physical activity. In addition, we used a novel renal marker (cystatin C) in contrast to creatinine $[30,51]$ to assess kidney function, and had extensive data on clinical characteristics that were considered potential confounders. However, our study has several limitations. First, due to the crosssectional design, we were unable to determine the causal nature of the associations observed. Second, due to the low response rate in our study, the results may be biased due to the voluntary participation of individuals who
had a high level of health consciousness and physical activity [52]. Third, we did not have data on the use of anti-platelet drugs. However, some participants on antiplatelet therapy may have been excluded to some extent because of the exclusion of those with a history of myocardial infarction or stroke. Fourth, the outcome variable, serum cystatin C levels, was measured once only, and we could not assess urinary albumin excretion. In addition, our study is limited by the small number of participants with low eGFR ( $<60 \mathrm{~mL} / \mathrm{min} / 1.73 \mathrm{~m}^{2}$ ). Finally, our study participants comprised residents of a rural community, where some of residents are engaged in fishing and farming. Since some studies suggest that physical activity differs between rural and urban residents [53, 54], caution is necessary when extrapolating our findings to people in urban areas, as well as general populations.

## Conclusions

Our data demonstrated an inverse association between sedentary behavior and kidney function in women, and a positive association between MVPA time and kidney function in both men and women. Future large-scale longitudinal studies are needed to determine the causal relationship between specific physical activity times and kidney dysfunction, which may help provide a clinical recommendation for various physical activity intensities.

## Abbreviations

BMI: Body mass index; CKD: Chronic kidney disease; CI: Confidence interval; DOSANCO: Dynamics of Lifestyle and Neighborhood Community; eGFR: Estimated glomerular filtration rate; HbA1c: Hemoglobin A1c; HDL: High-density lipoprotein; JSN-CKDI: Japan Society of Nephrology Chronic Kidney Disease Initiative; LPA: Light physical activity; METs: Metabolic equivalents; MVPA: Moderate-to-vigorous physical activity.

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## Authors' contributions

Research idea and study design: AT (Akiko Tamakoshi), KN, SU, EO, TN, AI, SS; data acquisition: KN, SS, SU, EO; data analysis/interpretation: KN, SS, SA, SI, TK, AY, AT (Aya Tanaka); statistical analysis: KN, SS; supervision or mentorship: AT (Akiko Tamakoshi), KN. All authors read and approved the final manuscript.

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## Availability of data and materials

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

## Declarations

## Ethics approval and consent to participate

All procedures performed in this study involving human participants were in accordance with the ethical standards of the institutional research committee and with the Declaration of Helsinki.

This study was approved by the institutional ethical board for epidemiological studies of Hokkaido University Graduate School of Medicine, approval number [18-018]. Written informed consent was obtained from all participants.

## Consent for publication

Not applicable.

## Competing interests

The authors declare that they have no competing interests.

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## References

1. Hill NR, Fatoba ST, Oke JL, Hirst JA, O'Callaghan CA, Lasserson DS, et al. Global prevalence of chronic kidney disease - a systematic review and Meta-analysis. PLoS One. 2016;11 (7):e0158765.
2. Imai E, Horio M, Watanabe T, Iseki K, Yamagata K, Hara S, et al. Prevalence of chronic kidney disease in the Japanese general population. Clin Exp Nephrol. 2009;13(6):621-30.
3. Go AS, Chertow GM, Fan D, McCulloch CE, Hsu CY. Chronic kidney disease and the risks of death, cardiovascular events, and hospitalization. N Engl J Med. 2004;351(13):1296-305.
4. Nagata M, Ninomiya T, Kiyohara Y, Murakami Y, Irie F, Sairenchi T, et al. Prediction of cardiovascular disease mortality by proteinuria and reduced kidney function: pooled analysis of 39,000 individuals from 7 cohort studies in Japan. Am J Epidemiol. 2013;178(1):1-11.
5. Matsushita K, Coresh J, Sang Y, Chalmers J, Fox C, Guallar E, et al. Estimated glomerular filtration rate and albuminuria for prediction of cardiovascular outcomes: a collaborative meta-analysis of individual participant data. Lancet Diabetes Endocrinol. 2015;3(7):514-25.
6. Whaley-Connell A, Sowers JR, McCullough PA, Roberts T, McFarlane SI, Chen SC, et al. Diabetes mellitus and CKD awareness: the kidney early evaluation program (KEEP) and National Health and nutrition examination survey (NHANES). Am J Kidney Dis. 2009;53(4 Suppl 4):S11-21.
7. Lee IM, Shiroma EJ, Lobelo F, Puska P, Blair SN, Katzmarzyk PT, et al. Effect of physical inactivity on major non-communicable diseases worldwide: an analysis of burden of disease and life expectancy. Lancet. 2012;380(9838):219-29.
8. Patterson R, McNamara E, Tainio M, de Sa TH, Smith AD, Sharp SJ, et al. Sedentary behaviour and risk of all-cause, cardiovascular and cancer mortality, and incident type 2 diabetes: a systematic review and dose response meta-analysis. Eur J Epidemiol. 2018;33(9):811-29.
9. Wilmot EG, Edwardson CL, Achana FA, Davies MJ, Gorely T, Gray LJ, et al. Sedentary time in adults and the association with diabetes, cardiovascular disease and death: systematic review and meta-analysis. Diabetologia. 2012;55(11):2895-905.
10. Hawkins MS, Sevick MA, Richardson CR, Fried LF, Arena VC, Kriska AM. Association between physical activity and kidney function: National Health and nutrition examination survey. Med Sci Sports Exerc. 2011;43(8):1457-64
11. Bharakhada N, Yates T, Davies MJ, Wilmot EG, Edwardson C, Henson J, et al. Association of sitting time and physical activity with CKD: a crosssectional study in family practices. Am J Kidney Dis. 2012;60(4):583-90.
12. Lee S, Shimada H, Lee S, Makizako H, Doi T, Harada K, et al. Association between sedentary time and kidney function in community-dwelling elderly Japanese people. Geriatr Gerontol Int. 2017;17(5):730-6.
13. Sallis JF, Saelens BE. Assessment of physical activity by self-report: status, limitations, and future directions. Res Q Exerc Sport. 2000;71 (2 Suppl):S1-14.
14. Hawkins MS, Storti KL, Richardson CR, King WC, Strath SJ, Holleman RG, et al. Objectively measured physical activity of USA adults by sex, age, and racial/ethnic groups: a cross-sectional study. Int J Behav Nutrition Phys Activity. 2009;6:31.
15. Glavinovic T, Ferguson T, Komenda P, Rigatto C, Duhamel TA, Tangri N, et al. CKD and sedentary time: results from the Canadian health measures survey. Am J Kidney Dis. 2018;72(4):529-37.
16. Gijsberts CM, Seneviratna A, de Carvalho LP, den Ruijter HM, Vidanapthirana $P$, Sorokin $V$, et al. Ethnicity modifies associations between cardiovascular risk factors and disease severity in parallel Dutch and Singapore coronary cohorts. PLoS One. 2015;10(7):e0132278.
17. Nakamura K, Hui SP, Ukawa S, Okada E, Nakagawa T, Okabe H, et al. Serum 25-hydroxyvitamin $D(3)$ levels and poor sleep quality in a Japanese population: the DOSANCO health study. Sleep Med. 2019;57:135-40.
18. Masse LC, Fuemmeler BF, Anderson CB, Matthews CE, Trost SG, Catellier DJ, et al. Accelerometer data reduction: a comparison of four reduction algorithms on select outcome variables. Med Sci Sports Exerc. 2005;37(11 Suppl):S544-54.
19. Oshima Y, Kawaguchi K, Tanaka S, Ohkawara K, Hikihara Y, Ishikawa-Takata K, Tabata I: Classifying household and locomotive activities using a triaxial accelerometer. In: Gait Posture. Volume 31, England: 2010 Elsevier B.V; 2010: 370-374.
20. Ohkawara K, Oshima Y, Hikihara Y, Ishikawa-Takata K, Tabata I, Tanaka S. Real-time estimation of daily physical activity intensity by a triaxial accelerometer and a gravity-removal classification algorithm. Br J Nutr. 2011;105(11):1681-91.
21. Kurita S, Yano S, Ishii K, Shibata A, Sasai H, Nakata Y, et al. Comparability of activity monitors used in Asian and Western-country studies for assessing free-living sedentary behaviour. PLoS One. 2017;12(10):e0186523.
22. Murakami H, Kawakami R, Nakae S, Yamada Y, Nakata Y, Ohkawara K, et al. Accuracy of 12 wearable devices for estimating physical activity energy expenditure using a metabolic chamber and the doubly labeled water method: validation study. JMIR Mhealth Uhealth. 2019;7(8):e13938.
23. Tudor-Locke C, Camhi SM, Troiano RP. A catalog of rules, variables, and definitions applied to accelerometer data in the National Health and nutrition examination survey, 2003-2006. Prev Chronic Dis. 2012;9:E113.
24. Migueles JH, Cadenas-Sanchez C, Ekelund U, Delisle Nyström C, MoraGonzalez J, Löf M, et al. Accelerometer Data Collection and Processing Criteria to Assess Physical Activity and Other Outcomes: A Systematic Review and Practical Considerations. Sports medicine (Auckland, NZ). 2017;47(9):1821-45.
25. Amagasa S, Inoue S, Ukawa S, Sasaki S, Nakamura K, Yoshimura A, et al. Are Japanese women less physically active than men? Findings from the DOSANCO health study. J Epidemiology / Japan Epidemiological Association. 2020.
26. Troiano RP, Berrigan D, Dodd KW, Masse LC, Tilert T, McDowell M. Physical activity in the United States measured by accelerometer. Med Sci Sports Exerc. 2008;40(1):181-8.
27. Yoshida J, Tsuji S, Tanaka T. New IT solution using human-oriented big data analytics. Hitachi Review. 2014;63(1):13-7.
28. Owen N, Healy GN, Matthews CE, Dunstan DW. Too much sitting: the population health science of sedentary behavior. Exerc Sport Sci Rev. 2010;38(3):105-13.
29. Horio M, Imai E, Yasuda Y, Watanabe T, Matsuo S. Collaborators developing the Japanese equation for estimated GFR: GFR estimation using standardized serum cystatin C in Japan. Am J Kidney Dis. 2013;61(2):197-203.
30. Oterdoom LH, Gansevoort RT, Schouten JP, de Jong PE, Gans RO, Bakker SJ. Urinary creatinine excretion, an indirect measure of muscle mass, is an independent predictor of cardiovascular disease and mortality in the general population. Atherosclerosis. 2009;207(2):534-40.
31. Umemura S, Arima H, Arima S, Asayama K, Dohi Y, Hirooka Y, et al. The Japanese Society of Hypertension Guidelines for the Management of Hypertension (JSH 2019). Hypertens Res. 2019;42(9):1235-481.
32. Haneda M, Noda M, Origasa H, Noto H, Yabe D, Fujita Y, et al. Japanese clinical practice guideline for diabetes 2016. Diabetol Int. 2018;9(1):1-45.
33. Kinoshita M, Yokote K, Arai H, lida M, Ishigaki Y, Ishibashi S, et al. Japan atherosclerosis society (JAS) guidelines for prevention of atherosclerotic cardiovascular diseases 2017. J Atheroscler Thromb. 2018;25(9):846-984
34. Kobayashi S, Murakami K, Sasaki S, Okubo H, Hirota N, Notsu A, et al Comparison of relative validity of food group intakes estimated by comprehensive and brief-type self-administered diet history questionnaires against 16 d dietary records in Japanese adults. Public Health Nutr. 2011;14(7):1200-11.
35. Jefferis BJ, Parsons TJ, Sartini C, Ash S, Lennon LT, Papacosta O, et al. Objectively measured physical activity, sedentary behaviour and all-cause mortality in older men: does volume of activity matter more than pattern of accumulation? Br J Sports Med. 2019;53(16):1013-20.
36. Parsons TJ, Sartini C, Ash S, Lennon LT, Wannamethee SG, Lee IM, et al. Objectively measured physical activity and kidney function in older men; a cross-sectional population-based study. Age Ageing. 2017;46(6):1010-4.
37. Lee J, Walker ME, Gabriel KP, Vasan RS, Xanthakis V. Associations of accelerometer-measured physical activity and sedentary time with chronic kidney disease: the Framingham heart study. PLoS One 2020;15(6):e0234825.
38. Levey AS, de Jong PE, Coresh J, El Nahas M, Astor BC, Matsushita K, et al. The definition, classification, and prognosis of chronic kidney disease: a KDIGO controversies conference report. Kidney Int. 2011;80(1):17-28.
39. Parvathaneni K, Surapaneni A, Ballew SH, Palta P, Rebholz CM, Selvin E et al. Association between midlife physical activity and incident kidney disease: the atherosclerosis risk in communities (ARIC) study. Am J Kidney Dis. 2021;77(1):74-81.
40. Borgundvaag E, Janssen I. Objectively measured physical activity and mortality risk among American adults. Am J Prev Med. 2017;52(1):e25-31.
41. Vaccarezza M, Papa V, Milani D, Gonelli A, Secchiero P, Zauli G, et al. Sex/Gender-Specific Imbalance in CVD: Could Physical Activity Help to Improve Clinical Outcome Targeting CVD Molecular Mechanisms in Women? Int J Mol Sci. 2020;21:4.
42. Bergens O, Nilsson A, Papaioannou KG, Kadi F. Sedentary patterns and systemic inflammation: sex-specific links in older adults. Front Physiol. 2021;12:625950.
43. Beddhu S, Baird BC, Zitterkoph J, Neilson J, Greene T. Physical activity and mortality in chronic kidney disease (NHANES III). Clin J Am Soc Nephrol. 2009;4(12):1901-6.
44. Johansen KL, Chertow GM, Ng AV, Mulligan K, Carey S, Schoenfeld PY, et al. Physical activity levels in patients on hemodialysis and healthy sedentary controls. Kidney Int. 2000;57(6):2564-70.
45. Wilhelm-Leen ER, Hall YN. M KT, Chertow GM: frailty and chronic kidney disease: the third National Health and nutrition evaluation survey. Am J Med. 2009;122(7):664-71 e662.
46. Moon SJ, Kim TH, Yoon SY, Chung JH, Hwang HJ. Relationship between stage of chronic kidney disease and sarcopenia in Korean aged 40 years and older using the Korea National Health and nutrition examination surveys (KNHANES IV-2, 3, and V-1, 2), 2008-2011. PLoS One. 2015;10(6):e0130740.
47. Izumi A, Kitamura M, Izawa KP. Effects of exercise training on delaying disease progression in patients with chronic kidney disease: a review of the literature. Rev Recent Clin Trials. 2016;11(4):333-41
48. Lee Y, Kwon S, Moon JJ, Han K, Paik NJ, Kim WS. The Effect of HealthRelated Behaviors on Disease Progression and Mortality in Early Stages of Chronic Kidney Disease: A Korean Nationwide Population-Based Study. J Clin Med. 2019:8(8).
49. Linke A, Erbs S, Hambrecht R. Effects of exercise training upon endothelial function in patients with cardiovascular disease. Front Biosci. 2008;13:424-32.
50. Katzmarzyk PT. Standing and mortality in a prospective cohort of Canadian adults. Med Sci Sports Exerc. 2014;46(5):940-6.
51. Shlipak MG, Matsushita K, Arnlov J, Inker LA, Katz R, Polkinghorne KR, et al. Cystatin $C$ versus creatinine in determining risk based on kidney function. N Engl J Med. 2013;369(10):932-43.
52. Inoue S, Ohya Y, Odagiri Y, Takamiya T, Kamada M, Okada S, et al. Characteristics of accelerometry respondents to a mail-based surveillance
study. Journal of epidemiology / Japan Epidemiological Association. 2010;20(6):446-52.
53. Klann A, Vu L, Ewing M, Fenton M, Pojednic R. Translating Urban Walkability Initiatives for Older Adults in Rural and Under-Resourced Communities. Int J Environ Res Public Health. 2019:16(17).
54. Fan JX, Wen M, Kowaleski-Jones L. Rural-urban differences in objective and subjective measures of physical activity: findings from the National Health and nutrition examination survey (NHANES) 2003-2006. Prev Chronic Dis. 2014;11:E141.

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