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# Retinal Vessel Diameter and Chronic Kidney Disease in Rural China

A Cross-Sectional Study

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**Abstract:** Few studies have examined the relationship between retinal microvascular abnormalities and chronic kidney disease (CKD). This study aims to examine the association between retinal vessel diameters and CKD in the rural China in order to provide the scientific basis for the early detection and diagnosis for CKD.

Participants and data were extracted from the Handan Eye Study, a population-based cross-sectional study performed from 2006 to 2007. The central retinal arteriolar equivalent (CRAE) and central retinal venular equivalent (CRVE) were summarized by the average arteriolar and venular caliber of each eye. The estimated glomerular filtration rate (eGFR) and a urinary albumin to creatinine ratio (ACR) were recorded. Multivariate logistic regression models were used to determine any associations between CRAE, CRVE, arteriole-to-venule ratio (AVR), retinopathy, and CKD in the recruited participants.

CKD was found in was 17.3% (892/5158) of this population with a 0.9% (48/5545) rate of reduced renal function and 16.7% (922/5538) rate of albuminuria. Retinopathy was present in 9.6% (571/5925) of participants. Compared to the 4th quartile of AVR, the first group was found to have a higher risk of albuminuria (odds ratio [OR] = 1.261, 95% confidence interval [95%CI]: 1.015–1.567, P = 0.037) and CKD (OR = 1.240, 95%CI: 1.000–1.537, P = 0.049) after adjustment for potential confounding variables. Retinopathy was associated with the occurrence of albuminuria (OR = 1.340, 95%CI: 1.067–1.685, P = 0.012) and CKD (OR = 1.341, 95%CI: 1.071–1.681, P = 0.010). In participants with diabetes, the ORs for the 1st and 4th quartiles of

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CRAE and CRVE were 2.292 (95%CI: 1.076–4.881, P=0.032) and 2.113 (95%CI: 1.006–4.438, P=0.048), respectively. Among the participants with hypertension, retinopathy was also observed to be associated with CKD (OR = 1.306, 95%CI: 1.003–1.699, P=0.047).

The parameters of retinal vessel diameter may be a useful index evaluating the occurrence and development of CKD.

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**Abbreviations**: ACR = albumin to creatinine ratio, AVR = arteriole-to-venule ratio, BMI = body mass index, CI = confidence interval, CKD = chronic kidney disease, CRAE = central retinal arteriolar equivalent, CRVE = central retinal venular equivalent, eGFR = estimated glomerular filtration rate, HDL = high-density lipoprotein, LDL = low-density lipoprotein, OR = odds ratio, SD = standard deviations, SiMES = Singapore Malay Eye Study.

## **INTRODUCTION**

C hronic kidney disease (CKD) has become a major public health problem worldwide and has been associated with premature morbidity and mortality.<sup>1–3</sup> CKD has long been a clinically challenging due to the late appearance of its clinical symptoms and the initiation of standard therapies only in late stages of the disease. Therefore, early detection, diagnosis, and treatment are the cornerstones of CKD management.

Some epidemiological studies have shown a relationship between CKD retinal microvascular abnormalities. Risk factors for microvascular damage, such as diabetes and hypertension, have been strongly associated with CKD,<sup>4–7</sup> and microvascular alterations in renal circulation in CKD have been documented in animal models.<sup>8</sup> Thus, microvascular disease has been suggested to play a major role in the etiology and evolution of CKD.

However, epidemiological studies are generally focused on urban and economically developed areas, while little is known about rural areas where lifestyle, level of education, cultural customs, nutrition, and environmental factors are very different.<sup>9–11</sup> Since rural population accounts for the vast majority of China's total populace, exploring the relationship between retinal microvascular abnormalities and CKD in this population is of great significance.

This study examines the association between retinal vessel diameters and CKD in rural China in order to provide a scientific basis for the early detection and diagnosis for CKD.

## MATERIALS AND METHODS

## **Study Population**

All patient data were extracted from the Handan Eye Study, a population-based cross-sectional study performed from 2006

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to 2007. A detailed explanation of this study and its methodology was published by Liang et al in 2009.<sup>9</sup> Residents of Yongnian County in Handan City, Hebei Province aged 6–30 years were randomly selected using a stratified, clustered, and multistaged sampling technique. Of the 458 villages in this region, 13 were randomly selected and stratified by geographic landform (plains or hillside). The protocol for this study was reviewed and approved by the Ethics Committee of the Beijing Tongren Hospital in accordance with the guidelines of the Helsinki Declaration, and written informed consent was obtained from all participants. Participants with missing information on serum creatinine, urinary albumin, or unclear retinal photography were excluded.

## Data Collection

The information collected from participants included sociodemographic status (age, sex, and education), past medical history (especially history of hypertension and/or diabetes), and lifestyle information (including assessment of smoking and drinking). Physical examinations were performed to measure height, weight, and blood pressure. Blood samples were obtained and fasting blood glucose, lipid levels, urea nitrogen, urea albumin, and creatinine were measured.<sup>9</sup> Albuminuria was measured by immunoturbidimetric methods (Audit Diagnostic, Ireland), and urinary creatinine was measured by the Jaffé kinetic method.

#### **Retinal Vessel Diameter Measurement**

The procedures for retinal photography and the grading of retinal microvascular signs have been described in a precursor study.<sup>9</sup> Retinal vessel diameters from digital retinal images were measured using a computer-based program by trained graders who were blinded to the status of each participant. The central retinal arteriolar equivalent (CRAE) and central retinal venular equivalent (CRVE) were summarized using the average arteriolar and venular caliber of each eye. This data were obtained by measuring all arterioles and venules coursing through a zone between 0.5 and 1 disc diameters away from the margin of the optic disc.<sup>10</sup>

Retinopathy was considered present if any characteristic lesion, as dened by the ETDRS severity scale, was present including microaneurysms, hemorrhages, cotton wool spots, intraretinal microvascular abnormalities, hard exudates, venous beading, and/or new vessels.

#### Definition of CKD

CKD was dened as the presence of either reduced renal function as evaluated by estimated glomerular filtration rate (eGFR, <60 mL/minute/1.73 m<sup>2</sup>) or albuminuria evaluated by urinary albumin to creatinine ratio (ACR,  $\geq$ 30 mg/g).<sup>11</sup> The eGFR was calculated using a modified version of the Modification of Diet in Renal Disease equation developed by the Peking University First Hospital, based on data from Chinese patients with CKD.<sup>12</sup>

## **Definition of Other Variables**

Hypertension was defined as systolic blood pressure  $\geq$ 140 mmHg and diastolic blood pressure  $\geq$ 90 mmHg, use of antihypertensive medication, or a self-reported previously diagnosed hypertension. Diabetes mellitus was defined as fasting blood glucose  $\geq$ 7.0 mmol/L, a self-reported history of diabetes, or those currently receiving insulin or oral hypoglycemic agents.

## Statistical Analysis

Body mass index (BMI) was calculated  $(kg/m^2)$  and participants were classified into either an overweight/obesity group ( $\geq 24.0 \text{ kg/m}^2$ ) or a normal/underweight group ( $< 24.0 \text{ kg/m}^2$ ). Level of education was based on number of years of schooling and classified into 2 groups,  $\geq 9$  and < 9 years. The arteriole-to-venule ratio (AVR) was calculated as the ratio of CRAE to CRVE. The parameters of CRAE, CRVE, and AVR were categorized into quartiles.

Population characteristics were described using means and standard deviations (SDs) or frequencies and proportions. The characteristics of the different quartiles of CRAE, CRVE, and AVR were also summarized and tested with one-way analysis of variance or the Chi-square test where appropriate. Multivariate logistic regression models were used to determine if associations existed between CRAE, CRVE, AVR, and/or retinopathy and CKD in the recruited population. Separate analyses were also performed for participants with diabetes, hypertension, and obesity. The effects of potential confounding variables and bias were controlled by adjusting the age, sex, BMI, education, hypertension, diabetes, smoking, drinking, total cholesterol, triglyceride, low-density lipoprotein (LDL), and high-density lipoprotein (HDL) levels.

Odds ratios (ORs) and their 95% confidence intervals (95%CIs) were estimated using the maximum likelihood method. A 2-sided *P* value of <0.05 was considered to be statistically significant. All analyses were performed using SAS software, version 9.1.3 (SAS Institute, Cary, NC).

#### RESULTS

A total of 5925 participants were recruited into the study. Retinal imaging and retinal vessel diameter measurements were obtained for all participants. eGFR data were available for 5158 participants and albuminuria data for 5538.

In this population, CKD had a prevalence of 17.3% (892/ 5158), reduced renal function in 0.9% (48/5545), albuminuria in 16.7% (922/5538), and retinopathy in 9.6% (571/5925) of participants. Basic characteristics of the participants are shown in Table 1. Participants with CKD were mostly older females and had greater incidences of hypertension, diabetes, and smoking (P < 0.05). Total cholesterol, triglycerides, and LDL levels were also found to be higher than in the general population (P < 0.05).

The differences between the CRAE quartiles were statistically significant for age, BMI, hypertension, diabetes, smoking, total cholesterol, LDL levels, and HDL levels (P < 0.05, Table 2). Sex, BMI, diabetes, smoking, alcohol consumption total cholesterol, triglycerides, LDL levels, and HDL levels revealed significant differences in the CRVE quartiles (P < 0.05, Table 3). The proportions of obesity, hypertension, diabetes, and mean total cholesterol, triglyceride, and LDL levels decreased with the increased AVR quartiles (P < 0.05, Table 4).

The associations between CRAE, CRVE, AVR, and CKD were analyzed with multivariate logistic regression models (Table 5). Compared to the 4th AVR quartile, the first quartile had a higher risk of albuminuria (OR = 1.261, 95%CI: 1.015–1.567, P = 0.037) and CKD (OR = 1.240, 95%CI: 1.000–1.537, P = 0.049) when adjusted by age, sex, incidence of smoking, alcohol consumption, BMI, level of education, and total cholesterol, triglycerides, and LDL, and HDL levels. Retinopathy was also associated with the occurrence of albuminuria (OR = 1.340, 95%CI: 1.067–1.685, P = 0.012) and CKD (OR = 1.341, 95%CI: 1.071–1.681, P = 0.010).

	Parameters Value	Ch	ronic Kidney Disease <sup>*</sup>		
Characteristics	(n = 5925)	Yes (n = 892)	No (n = 4266)	Р	
Age, years, mean $\pm$ SD	59.1±11.7	$55.3 \pm 11.5$	$51.1 \pm 11.3$	< 0.001	
Sex, female, %	3240 (54.7)	545 (61.1)	2289 (53.7)	< 0.001	
Education years $\geq 9, \%$	189 (3.2)	25 (2.8)	139 (3.3)	0.481	
BMI $\geq 24 \text{ kg/m}^2$ , %	2988 (50.4)	445 (49.9)	2191 (51.4)	0.424	
Hypertension	2874 (48.5)	555 (62.2)	1975 (46.3)	< 0.001	
Diabetes	376 (6.4)	84 (9.4)	238 (5.6)	< 0.001	
Smoking	1565 (26.4)	199 (22.3)	1155 (27.1)	0.003	
Drinking	1089 (18.4)	145 (16.3)	793 (18.6)	0.100	
Total cholesterol, mmol/L	$51.94 \pm 11.67$	$4.78 \pm 1.03$	$4.56\pm0.94$	< 0.001	
Triglyceride, mmol/L	$4.60\pm0.96$	$1.67 \pm 1.22$	$1.48 \pm 1.00$	< 0.001	
Low density lipoprotein, mmol/L	$1.51 \pm 1.05$	$2.82\pm0.68$	$2.68\pm0.64$	< 0.001	
High density lipoprotein, mmol/L	$1.28\pm0.29$	$1.29 \pm 0.30$	$1.27\pm0.29$	0.223	
Retinopathy, %	571 (9.6)	125 (14.0)	375 (8.8)	< 0.001	

### TABLE 1. The Basic Characteristics of the Participants Recruited in the Study

ACR = urinary albumin to creatinine ratio, eGFR = estimated glomerular filtration rate, SD = standard deviation.

\* Defined as the presence of reduced renal function and/or albuminuria among the 5158 participants with the data of eGFR and ACR.

In participants with diabetes, the ORs for CKD in the first CRAE and CRVE quartile 2.292 (95%CI: 1.076–4.881, P = 0.032) compared with 2.113 (95%CI: 1.006–4.438, P = 0.048) in the 4th quartile (Table 6). Among participants with hypertension, retinopathy was also found to be associated with CKD (OR = 1.306. 95%CI: 1.003–1.699, P = 0.047), as shown in Table 7. No significant differences were found between CRAE, CRVE, ACR, and CKD in participants with obesity (Table 8).

## DISCUSSION

Retinal microvasculature provides a unique window for noninvasive visualization of human circulation in vivo. Retinal microvasculature is considered as one of the best indicators of systemic microcirculation status in humans.<sup>3,6,7,13,14</sup> In recent years, thanks to the continuous development of fundus photography and digital image analysis technology, several large-scale epidemiological surveys have been conducted in other countries. These studies analyzed the relationship between retinal vascular abnormalities and changes in blood pressure, cardiovascular disease, kidney disease, and/or other systemic diseases.<sup>3,4,7,14–17</sup> Few studies reported a relationship between retinal vascular abnormalities and CKD, although the results were entirely comparable.<sup>6</sup>

According to the Atherosclerosis Risk in Communities Study, people with retinopathy were more likely to suffer from CKD (OR = 2.0; 95%CI: 1.4-2.8) after adjustment for potential confounding factors.<sup>6</sup> In the Singapore Malay Eye Study (SiMES), retinopathy was related to a decrease in eGFR and albuminuria.<sup>7</sup> In our analysis, we found a weaker association between retinopathy and CKD compared to the above studies.

In the Chronic Renal Insufficiency Cohort study, presence and severity of retinopathy at baseline were strongly associated

TABLE 2.	The Population	Characteristics	Grouped by	v C	Duartiles of	of	Central	Retinal	Arteriolar	Εαι	uivale	nt
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	Quartile of CRAE, µm								
Characteristics	Q1 (≤140.23)	Q2 (140.23-151.51)	Q3 (151.51-167.37)	Q4 (>167.37)	Р				
Age, years, mean $\pm$ SD	$53.53 \pm 12.06$	$51.90 \pm 11.19$	$51.23 \pm 11.73$	$51.06 \pm 11.52$	< 0.001				
Sex, female, %	803 (54.2)	820 (55.4)	805 (55.4)	812 (54.8)	0.918				
Education years $>9, \%$	43 (2.9)	46 (3.1)	50 (3.4)	50 (3.4)	0.858				
BMI $\geq 24 \text{ kg/m}^2, \%$	721 (48.7)	716 (48.4)	757 (51.1)	794 (53.6)	0.013				
Hypertension	935 (63.1)	726 (49.0)	623 (42.1)	590 (39.8)	< 0.001				
Diabetes	81 (5.5)	76 (5.1)	92 (6.2)	127 (8.6)	< 0.001				
Smoking	354 (23.9)	372 (25.1)	415 (28.0)	424 (28.6)	0.008				
Drinking	261 (17.6)	247 (16.7)	283 (19.1)	298 (20.1)	0.073				
Total cholesterol, mmol/L	$4.59\pm0.93$	$4.55\pm0.97$	$4.57\pm0.94$	$4.70\pm0.98$	< 0.001				
Triglyceride, mmol/L	$1.46\pm0.95$	$1.49 \pm 1.09$	$1.50 \pm 1.06$	$1.60 \pm 1.09$	0.003				
Low density lipoprotein, mmol/L	$1.26\pm0.26$	$1.26\pm0.28$	$1.27\pm0.28$	$1.31\pm0.32$	0.499				
High density lipoprotein, mmol/L	$2.71\pm0.64$	$2.70\pm0.67$	$2.69\pm0.64$	$2.73\pm0.66$	< 0.001				

CRAE = central retinal arteriolar equivalent, SD = standard deviation.

	Quartile of CRVE, µm							
Characteristics	Q1 (≤214.08)	Q2 (214.08-231.31)	Q3 (231.31-257.41)	Q4 (>257.41)	Р			
Age, years, mean $\pm$ SD	$52.19 \pm 12.19$	$51.63 \pm 11.50$	$52.3 \pm 11.64$	$51.62 \pm 11.34$	0.237			
Sex, female, %	850 (57.4)	778 (52.5)	822 (55.5)	790 (53.3)	0.035			
Education years $\geq 9, \%$	52 (3.5)	38 (2.6)	44 (3.0)	55 (3.7)	0.271			
BMI $\geq 24 \text{ kg/m}^2$ , %	677 (45.7)	705 (47.6)	767 (51.8)	839 (56.7)	< 0.001			
Hypertension	748 (50.5)	720 (48.6)	725 (48.9)	681 (46.0)	0.100			
Diabetes	72 (4.9)	66 (4.5)	100 (6.8)	138 (9.3)	< 0.001			
Smoking	328 (22.2)	422 (28.5)	388 (26.2)	427 (28.8)	< 0.001			
Drinking	237 (16.0)	288 (19.5)	269 (18.2)	295 (19.9)	0.028			
Total cholesterol, mmol/L	$4.51\pm0.93$	$4.50 \pm 0.93$	$4.65 \pm 0.95$	$4.75\pm0.99$	< 0.001			
Triglyceride, mmol/L	$1.39\pm0.94$	$1.46 \pm 1.05$	$1.56 \pm 1.10$	$1.64 \pm 1.08$	< 0.001			
Low density lipoprotein, mmol/L	$2.65\pm0.63$	$2.67 \pm 0.64$	$2.74 \pm 0.65$	$2.76\pm0.67$	< 0.001			
High density lipoprotein, mmol/L	$1.27\pm0.26$	$1.25\pm0.27$	$1.27\pm0.28$	$1.31\pm0.33$	< 0.001			

with the risk of subsequent progression to end-stage renal disease and reductions in eGFR in univariate analyses. However, this association was not statistically significant after adjustment for initial eGFR and 24-hour proteinuria.<sup>18</sup> Adjustment for all potential confounding factors is extremely important in order to determine the association between retinopathy and CKD, and further studies may be warranted.

In this study, we did not find an association between quartiles of CRAE and CRVE and CKD considering the whole sample or just hypertensive participants. The presence of retinal arteriolar abnormalities was also not associated with deteriorating renal function in the Cardiovascular Health Study.<sup>19</sup> On the other hand, in the Atherosclerosis Risk in Communities Study, patients with retinal arteriolar/venular narrowing were more likely to suffer from CKD compared to the general population. In the SiMES, only retinal arteriolar narrowing was related to increased albuminuria, while no difference was found for retinal venular changes.<sup>7</sup> Interestingly, we found that the likelihood of suffering from CKD increased with decrease in retinal diameter. Therefore, AVR may act as a marker for the early stages of CKD. However, this association could not be observed once the patients were stratified based on hypertension and diabetes diagnoses. This confirms the results obtained by Masaidi et  $al^{20}$  who reported that no significant associations could be found between AVR and eGFR/microalbuminuria in hypertensive patients. However, our results could be explained by the specificity of this population.

The association between CKD/albuminuria and retinopathy has been evaluated innumerousstudies.<sup>13,16,21–23</sup> For example, the SiMES showed that retinal arteriolar narrowing was associated with CKD.<sup>19</sup> We did not find statistically significant associations between these variables in the general population, but we did find that retinal arterial or venular narrowing increases as CKD progresses in patients with diabetes. Previous studies have also reported that type 2 diabetic

TABLE 4.	The Population	Characteristics	Grouped by	/ Quartiles	of Arteriole-to-Venule Ratio
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	Quartile of AVR							
Characteristics	Q1 (≤0.61)	Q2 (0.61-0.65)	Q3 (0.65-0.70)	Q4 (>0.70)	Р			
Age, years, mean $\pm$ SD	$54.33 \pm 11.46$	$51.49 \pm 11.44$	$50.63 \pm 11.37$	$51.22 \pm 12.17$	0.237			
Sex, female, %	828 (53.2)	793 (54.8)	907 (54.0)	712 (54.5)	0.128			
Education years $>9, \%$	53 (3.4)	54 (3.7)	42 (2.5)	40 (3.2)	0.242			
BMI $\geq 24 \text{ kg/m}^2$ , %	850 (54.6)	752 (51.9)	834 (49.6)	552 (44.6)	< 0.001			
Hypertension	992 (63.7)	711 (49.1)	708 (42.1)	463 (37.4)	< 0.001			
Diabetes	129 (8.3)	92 (6.4)	88 (5.2)	67 (5.4)	0.002			
Smoking	404 (25.9)	385 (26.6)	448 (26.7)	328 (26.5)	0.966			
Drinking	292 (18.7)	288 (19.5)	269 (18.2)	295 (19.9)	0.028			
Total cholesterol, mmol/L	$4.73 \pm 0.97$	$4.63 \pm 0.97$	$4.55 \pm 0.93$	$4.5 \pm 0.95$	< 0.001			
Triglyceride, mmol/L	$1.6 \pm 1.05$	$1.53 \pm 1.07$	$1.49 \pm 1.06$	$1.41 \pm 1.01$	< 0.001			
Low density lipoprotein, mmol/L	$2.8\pm0.66$	$2.72\pm0.67$	$2.67\pm0.63$	$2.63\pm0.62$	< 0.001			
High density lipoprotein, mmol/L	$1.27\pm0.29$	$1.27\pm0.29$	$1.28\pm0.28$	$1.28\pm0.28$	0.541			

AVR = arteriole-to-venule ratio, SD = standard deviation.

Ratio and Chronic Kidney Disease in All The Participants	TABLE 5.	The Associations	Among Central Retin	hal Arteriolar E	quivalent, C	Central Retinal	Venular Equivalent,	Arteriole-to-Venule
	Ratio and	Chronic Kidney E	Disease in All The Pa	rticipants	•		•	

	<b>Reduced Renal Function</b>				Albuminuria				
Variable	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р
CRAE									
Q1	10 (0.7)	0.747 (0.311, 1.795)	0.515	252 (18.1)	1.190 (0.962, 1.471)	0.109	236 (18.7)	1.169 (0.948, 1.442)	0.145
Q2	14 (1.0)	1.228 (0.544, 2.769)	0.621	227 (16.4)	1.112 (0.897, 1.378)	0.333	220 (17.4)	1.102 (0.891, 1.362)	0.370
Q3	12 (0.9)	1.109 (0.482, 2.552)	0.807	231 (16.7)	1.145 (0.925, 1.415)	0.213	223 (17.4)	1.117 (0.905, 1.379)	0.303
Q4	12 (0.8)	Reference		212 (15.4)	Reference		213 (15.8)	Reference	
CRVE									
Q1	14 (1.1)	0.994 (0.445, 2.219)	0.988	227 (16.2)	0.968 (0.780, 1.200)	0.766	208 (16.5)	0.964 (0.780, 1.193)	0.738
Q2	11 (0.8)	0.897 (0.387, 2.078)	0.800	227 (16.4)	1.047 (0.847, 1.294)	0.673	219 (17.4)	1.043 (0.846, 1.287)	0.693
Q3	10 (0.7)	0.668 (0.282, 1.582)	0.359	245 (17.7)	1.109 (0.902, 1.364)	0.327	237 (18.3)	1.079 (0.879, 1.325)	0.466
Q4	13 (0.9)	Reference		223 (16.3)	Reference		228 (17.0)	Reference	
AVR									
Q1	15 (1.0)	0.759 (0.355, 1.623)	0.477	286 (19.8)	1.261 (1.015, 1.567)	0.037	279 (20.7)	1.240 (1.000, 1.537)	0.049
Q2	11 (0.8)	0.769 (0.337, 1.754)	0.532	211 (15.7)	1.011 (0.805, 1.269)	0.927	208 (16.5)	1.016 (0.812, 1.272)	0.891
Q3	8 (0.5)	0.560 (0.228, 1.378)	0.207	240 (15.3)	1.021 (0.818, 1.273)	0.856	230 (15.7)	1.003 (0.806, 1.249)	0.978
Q4	14 (1.2)	Reference		185 (15.8)	Reference		175 (16.2)	Reference	
Retinopathy	10 (1.8)	1.431 (0.691, 2.960)	0.334	127 (24.1)	1.340 (1.067, 1.685)	0.012	125 (25.0)	1.341 (1.071, 1.681)	0.010

AVR = arteriole-to-venule ratio, CI = confidence interval, CRAE = central retinal arteriolar equivalent, CRVE = central retinal venular equivalent, OR = odds ratio.

patients with proliferative diabetic retinopathy are more likely to present with renal involvement.<sup>23</sup> Therefore, all patients with proliferative diabetic retinopathy or with retinal arterial or venular narrowing should undergo an evaluation of renal function, including urinary albumin measurements. Zhang et al<sup>24</sup> reported that the prevalence of CKD was

Zhang et al<sup>2+</sup> reported that the prevalence of CKD was 14.3% in 13 provinces in China – not including Hebei province analyzed herein – with 2.8% of the population having reduced renal function and 7.3% having albuminuria. Xue et al<sup>25</sup>

sampled residents of Guangxi Province and reported that the prevalence of albuminuria and reduced renal function were 12.5% and 0.4%, respectively – the overall CKD prevalence was 15.3%. Compared to these studies, we have found similar prevalence of CKD but a higher rate of albuminuria which may be related to lifestyle differences.

The main limitation of this study is its cross-sectional nature which limits determination of causal inferences. Secondly, albuminuria was measured only once using urinary ACR,

**TABLE 6.** The Associations Among Central Retinal Arteriolar Equivalent, Central Retinal Venular Equivalent, Arteriole-to-Venule Ratio and Chronic Kidney Disease in the Participants With Diabetes

	<b>Reduced Renal Function</b>				Albuminuria				
Variable	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р
CRAE									
Q1	3 (4.2)	4.052 (0.348, 47.178)	0.264	21 (28.8)	1.975 (0.914, 4.266)	0.083	23 (35.9)	2.292 (1.076, 4.881)	0.032
Q2	2 (2.8)	1.969 (0.136, 28.417)	0.619	22 (31.9)	2.172 (1.033, 4.568)	0.041	22 (34.4)	1.962 (0.936, 4.113)	0.074
Q3	3 (3.5)	4.669 (0.398, 54.723)	0.220	19 (23.5)	1.273 (0.591, 2.742)	0.538	18 (24.0)	1.264 (0.593, 2.693)	0.544
Q4	1 (0.8)	Reference		20 (16.8)	Reference		21 (17.7)	Reference	
CRVE									
Q1	3 (4.7)	2.846 (0.340, 23.826)	0.335	21 (32.3)	1.918 (0.907, 4.058)	0.088	22 (38.6)	2.113 (1.006, 4.438)	0.048
Q2	0	NA	0.955	17 (27.9)	1.365 (0.624, 2.986)	0.435	16 (29.1)	1.268 (0.578, 2.783)	0.553
Q3	4 (4.2)	1.968 (0.301, 12.862)	0.480	20 (21.7)	1.116 (0.552, 2.255)	0.76	21 (24.1)	1.179 (0.591, 2.354)	0.640
Q4	2 (1.5)	Reference		24 (19.4)	Reference		25 (20.3)	Reference	
AVR									
Q1	5 (4.1)	1.605 (0.144, 17.915)	0.701	35 (29.9)	1.083 (0.510, 2.301)	0.836	37 (33.6)	1.202 (0.565, 2.557)	0.633
Q2	0	NA	0.950	17 (21.0)	0.626 (0.272, 1.440)	0.270	16 (21.1)	0.604 (0.261, 1.400)	0.240
Q3	3 (3.5)	1.728 (0.142, 21.015)	0.668	12 (14.8)	0.421 (0.176, 1.007)	0.052	14 (18.0)	0.497 (0.212, 1.165)	0.108
Q4	1 (1.6)	Reference		18 (28.6)	Reference		17 (29.3)	Reference	
Retinopathy	4 (3.5)	1.636 (0.353, 7.583)	0.529	35 (32.4)	1.691 (0.972, 2.943)	0.063	37 (35.2)	1.625 (0.939, 2.810)	0.083

AVR = arteriole-to-venule ratio, CI = confidence interval, CRAE = central retinal arteriolar equivalent, CRVE = central retinal venular equivalent, OR = odds ratio.

	<b>Reduced Renal Function</b>				Albuminuria				
Variable	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р
CRAE									
Q1	9 (1.0)	0.655 (0.256, 1.677)	0.378	179 (20.4)	1.106 (0.837, 1.461)	0.480	177 (21.6)	1.077 (0.818, 1.416)	0.598
Q2	11 (1.6)	1.009 (0.410, 2.484)	0.985	141 (21.0)	1.132 (0.845, 1.518)	0.406	142 (22.7)	1.120 (0.839, 1.493)	0.442
Q3	11 (1.9)	1.172 (0.479, 2.868)	0.728	127 (22.1)	1.067 (0.790, 1.442)	0.670	121 (22.1)	1.040 (0.774, 1.398)	0.794
Q4	10 (1.7)	Reference		109 (20.0)	Reference		115 (21.5)	Reference	
CRVE									
Q1	12 (1.7)	0.789 (0.337, 1.851)	0.587	140 (20.0)	1.011 (0.761, 1.343)	0.939	137 (21.0)	0.992 (0.751, 1.311)	0.957
Q2	9 (1.3)	0.692 (0.279, 1.712)	0.426	137 (20.5)	1.075 (0.810, 1.428)	0.616	136 (21.8)	1.062 (0.804, 1.403)	0.673
Q3	8 (1.2)	0.481 (0.188, 1.229)	0.126	155 (23.0)	1.187 (0.901, 1.564)	0.223	151 (23.9)	1.123 (0.856, 1.474)	0.402
Q4	12 (1.8)	Reference		124 (19.8)	Reference		131 (21.2)	Reference	
AVR									
Q1	14 (1.5)	0.897 (0.383, 2.102)	0.803	206 (22.4)	1.150 (0.861, 1.537)	0.344	208 (23.9)	1.150 (0.864, 1.530)	0.339
Q2	9 (1.3)	0.844 (0.328, 2.169)	0.724	127 (19.3)	0.964 (0.705, 1.319)	0.818	129 (20.5)	0.979 (0.719, 1.332)	0.892
Q3	8 (1.2)	0.875 (0.330, 2.318)	0.789	128 (19.4)	0.968 (0.707, 1.327)	0.841	124 (20.0)	0.970 (0.711, 1.323)	0.846
Q4	10 (2.2)	Reference		95 (22.1)	Reference		94 (22.8)	Reference	
Retinopathy	10 (2.6)	1.618 (0.770, 3.399)	0.204	98 (26.6)	1.288 (0.986, 1.683)	0.064	98 (28.1)	1.306 (1.003, 1.699)	0.047

**TABLE 7.** The Associations Among Central Retinal Arteriolar Equivalent, Central Retinal Venular Equivalent, Arteriole-to-Venule Ratio and Chronic Kidney Disease in the Participants With Hypertension

AVR = arteriole-to-venule ratio, CI = confidence interval, CRAE = central retinal arteriolar equivalent, CRVE = central retinal venular equivalent, OR = odds ratio.

TABLE 8. The Associations Among Central Retinal Arteriolar Equivalent, Central Retinal Venular Equivalent, Arteriole-to-Venule Ratio and Chronic Kidney Disease in the Participants With Overweight/Obesity

	Reduced Renal Function				Albuminuria				
Variable	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р	n, %	OR (95%CI)	Р
CRAE									
Q1	2 (0.3)	0.288 (0.056, 1.472)	0.135	118 (17.4)	1.083 (0.809, 1.450)	0.591	111 (17.7)	1.056 (0.790, 1.411)	0.712
Q2	10 (1.5)	1.763 (0.614, 5.063)	0.292	97 (14.5)	0.893 (0.661, 1.206)	0.459	100 (16.1)	0.930 (0.692, 1.249)	0.630
Q3	5 (0.7)	0.910 (0.269, 3.082)	0.880	112 (16.1)	1.009 (0.755, 1.347)	0.954	108 (16.3)	0.980 (0.735, 1.307)	0.891
Q4	7 (0.9)	Reference		125 (16.9)	Reference		126 (17.4)	Reference	
CRVE									
Q1	6 (1.0)	1.642 (0.458, 5.890)	0.447	99 (15.4)	0.960 (0.710, 1.297)	0.789	93 (15.7)	0.972 (0.721, 1.311)	0.855
Q2	7 (1.1)	1.631 (0.474, 5.609)	0.438	101 (15.5)	0.974 (0.724, 1.312)	0.864	101 (16.8)	1.013 (0.756, 1.359)	0.929
Q3	6 (0.8)	1.449 (0.412, 5.096)	0.563	121 (16.8)	1.087 (0.822, 1.438)	0.557	120 (17.7)	1.080 (0.818, 1.426)	0.589
Q4	5 (0.6)	Reference		131 (16.9)	Reference		131 (17.2)	Reference	
AVR									
Q1	5 (0.6)	0.424 (0.128, 1.407)	0.161	148 (18.7)	1.282 (0.935, 1.757)	0.123	144 (19.3)	1.222 (0.895, 1.667)	0.207
Q2	6 (0.8)	0.702 (0.223, 2.212)	0.546	99 (14.2)	0.941 (0.672, 1.318)	0.724	98 (14.8)	0.926 (0.665, 1.290)	0.649
Q3	6 (0.8)	0.724 (0.225, 2.325)	0.587	127 (16.4)	1.180 (0.856, 1.626)	0.312	127 (17.3)	1.143 (0.833, 1.568)	0.407
Q4	7 (1.3)	Reference		78 (14.9)	Reference		76 (15.4)	Reference	
Retinopathy	5 (1.9)	1.658 (0.591, 4.656)	0.337	56 (22.5)	1.345 (0.965, 1.874)	0.080	57 (23.6)	1.352 (0.976, 1.874)	0.070

AVR = arteriole-to-venule ratio, CI = confidence interval, CRAE = central retinal arteriolar equivalent, CRVE = central retinal venular equivalent, OR = odds ratio.

which may overestimate or underestimate the real prevalence of albuminuria. Additionally, a selection bias may have occurred due to exclusion of eligible participants with incomplete records. The results from this population-based data from rural China suggest that retinal vessel diameter is associated with CKD, especially in the diabetic population, and may serve as a useful index for evaluating the occurrence and development of CKD.

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