**Open Access** Research

# BMJ Open Relationship among diet habit and lower urinary tract symptoms and sexual function in outpatient-based males with LUTS/BPH: a multiregional and cross-sectional study in China

Yuke Chen,<sup>1</sup> Wei Yu,<sup>1</sup> Liqun Zhou,<sup>1</sup> Shiliang Wu,<sup>1</sup> Yang Yang,<sup>1</sup> Jianye Wang,<sup>2</sup> Ye Tian,<sup>3</sup> Dalin He,<sup>4</sup> Yong Xu,<sup>5</sup> Jian Huang,<sup>6</sup> Xiaofeng Wang,<sup>7</sup> Xin Gao,<sup>8</sup> Hanzhong Li,<sup>9</sup> Lulin Ma,<sup>10</sup> Ning Zhang,<sup>11</sup> Shengtian Zhao,<sup>12</sup> Xunbo Jin<sup>13</sup>

To cite: Chen Y. Yu W. Zhou L. et al. Relationship among diet habit and lower urinary tract symptoms and sexual function in outpatientbased males with LUTS/BPH: a multiregional and crosssectional study in China. BMJ Open 2016;6:e010863. doi:10.1136/bmjopen-2015-010863

Prepublication history and additional material is available. To view please visit the journal (http://dx.doi.org/ 10.1136/bmjopen-2015-010863).

YC and WY contributed equally.

Received 17 December 2015 Revised 19 July 2016 Accepted 29 July 2016



For numbered affiliations see end of article.

#### Correspondence to

Professor Ligun Zhou: zhoulgmail@sina.com and Shiliang Wu; wushiliangjsh@263.net

#### **ABSTRACT**

**Objectives:** This study assessed the effect of diet habits on lower urinary tract symptoms (LUTS) and sexual function in Chinese men with LUTS/benign prostatic hypertrophy (LUTS/BPH).

Setting: Multicentre study conducted between July 2013 and December 2013 in 11 hospitals in 3 geographic regions in China.

Participants: Overall, participants with LUTS/BPH accounted for 61.4% (2584/4208) of the respondents, whose data were processed in the following statistical analysis.

### Primary and secondary outcome measures:

LUTS and sexual function were assessed based on the International Prostate Symptom Score (IPSS) and the International Index of Erectile Function 5 (IIEF-5) score. Prostate volume (PV) was determined by ultrasound.

Results: A total of 4208 participants met the inclusion criteria. The average age of the whole participants was 65.8±7.7 years. Overall, participants with LUTS/BPH accounted for 61.4% (2584/4208) of the respondents, whose data were processed in the following statistical analysis. Generally, prostate enlargement was greatest in south China. LUTS and male sexual dysfunction (MSD) were most severe in northwest China. Based on multivariable analysis, PV enlarged as the age (p<0.001), body mass index (BMI; p<0.001) and vegetable intake (p<0.001) increased. Age (p<0.001) and BMI (p<0.05) independently increased the IPSS. A higher level of education (p<0.001) and more frequent meat, fish and egg intake (p<0.05) decreased the IPSS. Age (p<0.001), BMI (p<0.001), low education level (p<0.05), vegetable intake (p=0.001), and milk and dairy product intake (p=0.001) decreased the IIEF-5 score.

**Conclusions:** In addition to factors including age, obesity and level of education, dietary habits and geographic difference might also play an important role in the variation of PV, LUTS and MSD for Chinese men with LUTS/BPH.

## Strengths and limitations of this study

- The first strength of the study is its relatively large sample size and multiregional design.
- The second strength of the study is attributed to the detection of geographic difference in diet habits and clinical outcomes (the prostate volume. International Prostate Symptom Score (IPSS) and International Index of Erectile Function 5 (IIEF-5) score), which might support the planning and implementation of public health policies.
- The interesting associations of prostate volume and IPSS under the influence of geography and diet contribute to another strength of the study.
- However, the limitations of the study include the regional sample size diversity, selection bias of participants and broad classification of diet.

#### INTRODUCTION

Benign prostatic hyperplasia (BPH) is very common among ageing men and causes lower urinary tract symptoms (LUTS), which lead to a diminished health-related quality of life. Although BPH was rare in some Asian countries in the early years of the 20th century, it has become a common disease in recent decades, the prevalence of which is comparable to the West.<sup>2</sup> Accumulating evidence suggests that other than ageing, modifiable factors, such as increasing prostate volume (PV), obesity, diet, dyslipidaemia, hormonal imbalance, hypertension, metabolic syndrome, alcohol and smoking, also contribute to the development of BPH, LUTS or both.  $^{1}$   $^{3-5}$ 

The prevalence of erectile dysfunction (ED) in elderly men has been reported to be 23.2% in Australia, 19.2% in Germany and

12% in China.<sup>2</sup> An increasing number of studies in recent years have suggested that patients with BPH or LUTS may be more prone to ED, and that LUTS/BPH symptoms often coexist with ED.<sup>8</sup> Additionally, age, household income, lifestyle, smoking and alcohol consumption, obesity, and physical inactivity have also been reported to be associated with the occurrence and severity of ED.<sup>10–12</sup> Recently, epidemiological data suggest the effects of nutritional factors modulating metabolism substantially prior to the development of BPH, LUTS and ED.<sup>13–15</sup>

In the current study, we used data from a recent outpatient-based screening survey to assess the epidemiological characteristics of prostate health and male sexual function in different regions in China.

#### PATIENTS AND METHODS

The data were collected between July 2013 and December 2013 in three geographic regions in China (Beijing and Tianjin (north China), Guangzhou (south China) and Xi'an (northwest China)). First, leaflets, printed with health education concerning prostate disease and information about a clinical check-up on the prostate, were handed out in the communities in the above three regions. Then our sample consisted of 5484 native males who attended the clinic for a prostate check-up at medical centres (The First, Second, and Third Affiliated Hospitals of Peking University, Peking Union Medical College Hospital, Beijing Chao-Yang Hospital, Beijing Friendship Hospital, Beijing Hospital in Beijing, The Second Affiliated Hospital of Tianjin Medical University in Tianjin, The Second and Third Affiliated Hospitals of Zhongshan University Guangzhou, and The First Affiliated Hospitals of Xi'an Jiaotong University in Xi'an). Temporary residents or refugees were not involved in this study. Participants who were <50 years of age were excluded. The exclusion criteria included urinary deformities, urinary injuries, urinary neoplasms, urethral strictures, neurobladders, urinary tract infections, acute prostatitis, treatment for BPH or sexual dysfunction, and treatments with known urinary tract or sexual function side effects. The study was approved by the Institutional Review Board at all participating centres. All participants provided written informed consent.

The research was carried out by trained interviewers at the respective centres. The interviewers were clinical physicians who were trained uniformly to be certified in conducting the survey. The main tasks for the interviewers were to deliver the survey, instruct participants on how to fill out the questionnaire, and collect the participants' information from completed questionnaires and clinical tests.

Peking University Clinical Research Institute was responsible for creating, upgrading, maintaining, backing up and recovering the database according to standard operating procedures, and for checking the data. The questionnaires were entered in the database twice by different individuals, and both versions were compared to ensure the accuracy of the data.

The structured questionnaire elicited information regarding age, level of education ( $\leq 9$  years or > 9 years), height, weight, tobacco and alcohol use (never/ever or current user), weekly common food intake frequency, medical history, recent drug history, LUTS conditions according to the International Prostate Symptom Score (IPSS) questionnaire, and sexual function conditions according to the International Index of Erectile Function-5 (IIEF-5). Body mass index (BMI) was calculated as the weight in kg/height in m<sup>2</sup>. Dietary intake over the past 1 year period was assessed and sorted into three categories (vegetables (green vegetables, carrot and tomato); meat (pork, beef and chicken), fish and eggs; and milk and dairy products). The monthly consumption of each dietary category was obtained from weekly consumption multiplied by 4, which was then distributed into tertiles based on the distribution of the entire study population and defined as low, moderate and high. 16 Additionally, the following clinical examinations were performed: serum prostate-specific antigen (PSA) level; routine urinalysis; prostatic fluid analysis; digital rectal examination (DRE) and transrectal ultrasound (TRUS). The type of TRUS used varied among the participating institutions, but all measurements were made using a 7.5 MHz rectal probe. PV was calculated by substituting the formula for an ellipsoid, as follows:  $\pi/6 \times (\text{height}) \times (\text{length}) \times (\text{width})$ , with the height, length and width of the prostate measured by TRUS.

A prostate biopsy was obtained from all participants with a PSA level >4 ng/mL or findings suspicious of prostate cancer on DRE. Those participants who were diagnosed with prostate cancer were excluded from the study.

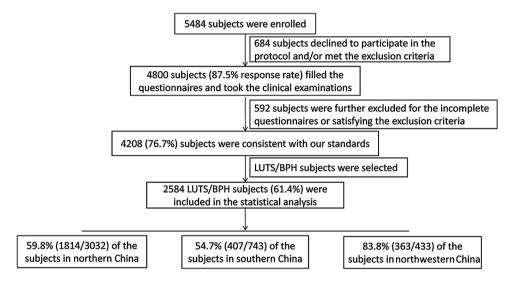
LUTS/BPH was defined as ≥50 years of age and an IPSS ≥8; LUTS/BPH data were used for the final analysis. SPSS V.16.0 was used for statistical analysis. The means and SDs were used to present normally distributed continuous variables; the median (q1 and q3) was used to present non-normally distributed variables. The Kruskal-Wallis test or one-way analysis of variance (ANOVA) was used to compare the dietary habits, PV, IPSS and IIEF-5 score of participants with LUTS/BPH from the three geographic regions. In the univariate part: the t-test or one-way ANOVA was used for PV and IPSS and the Wilcoxon test or Kruskal-Wallis test was used for the IIEF-5 score in categorical variables (education, smoking, alcohol consumption and region). Univariate linear regression was used for all the three clinical outcomes in continuous variables (age, BMI and diet). Multiple linear regression was performed as a multivariate analysis of the PV, IPSS and IIEF-5 score. The IIEF-5 score was log-changed to conform to a normal distribution. A dummy variable change was performed for the variable of the region to satisfy the multiple linear logistic model. Finally, univariate and multiple linear regression (adjusted for age, level of education, BMI, smoking and alcohol consumption, and diet) were performed to detect the associations of the PV, IPSS and IIEF-5 score in the entire cohort. A p<0.05 was considered statistically significant.

#### **RESULTS**

The process of the study design and participant exclusion is presented in figure 1. Of the 5484 enrolled participants, 4800 filled the questionnaire and underwent all of the clinical tests; 684 declined to participate in the protocol and/or met exclusion criteria. Thus, the response rate was 87.5%. Five hundred and ninety-two participants were further excluded for the incomplete questionnaires or satisfying the exclusion criteria. The data from 4208 (76.7%) participants were consistent with our standards. The average age of the 4208 participants was 65.8±7.7 years. The overall prevalence of

LUTS/BPH among the study participants was 61.4% (2584/4208); 59.8% (1814/3032), 54.7% (407/743) and 83.8% (363/433) of the participants resided in northern China, southern China and northwestern China, respectively (table 1). Therefore, the data from 2584 participants were included in the statistical analysis.

The dietary and clinical characteristics (the PV, IPSS and IIEF-5 score as a function of different age groups) among the participants with LUTS/BPH residing in the three geographic regions are listed in table 2. Participants in southern China consumed the highest amounts of meat, fish and eggs, while participants in northern China and northwestern China consumed the highest amounts of vegetables and milk and dairy products, respectively. The PV, IPSS and IIEF-5 score followed a deteriorating trend with ageing. PV was significantly different among the participants residing in the three geographic regions for all age groups (p<0.01). It was largest among men residing in southern



**Figure 1** Flow diagram for explaining our study design and participants excluded from analysis. LUTS/BPH was defined as an age ≥50 years and an IPSS≥8. IPSS, International Prostate Symptom Score; LUTS/BPH, lower urinary tract symptoms/ benign prostatic hypertrophy.

Characteristics	Northern China (n=3032)	Southern China (n=743)	Northwestern China (n=433)	Total (n=4208)
LUTS/BPH, n (%)	1814 (59.8)	407 (54.7)	363 (83.8)	2584 (61.4)
Age (years) (means±SDs)	66.1±8.0	65.5±8.3	65.1±7.7	65.8±7.7
BMI (kg/m <sup>2</sup> ) (means±SDs)	24.6±4.5	23.6±3.2	24.1±2.5	24.3±4.1
Education (years), n (%)				
≤9	690 (22.8)	193 (26.0)	129 (29.8)	1012 (24.0)
>9	2342 (77.2)	550 (74.0)	304 (70.2)	3196 (76.0)
Smoking, n (%)				
No	1835 (60.7)	399 (53.7)	257 (59.4)	2491 (59.2)
Yes	1197 (39.3)	344 (46.3)	176 (40.6)	2717 (40.8)
Alcohol consumption, n (%)				
No	1910 (63.0)	459 (61.8)	279 (64.4)	2648 (62.9)
Yes	1122 (37.0)	284 (38.2)	154 (35.6)	1560 (37.1)

Table 2 Comparison of dietary (stratified by tertile) and clinical characteristics (stratified by age decades) among LUTS/BPH participants from the three geographic regions

Characteristics	Northern China (n=1814)	Southern China (n=407)	Northwestern China (n=363)	p Value
Dietary (monthly)				
Meat, fish and eggs, n (%)				
Low (<16 times/month)	347 (19.1)	7 (1.7)	82 (22.6)	
Moderate (16–28 times/month)	503 (27.7)	86 (21.1)	242 (66.7)	
High (>28 times/month)	964 (53.1)	314 (77.1)	39 (10.7)	< 0.001
Vegetables				
Low (<24 times/month)	606 (33.4)	134 (32.9)	121 (33.3)	
Moderate (24–32 times/month)	542 (29.9)	180 (44.2)	240 (66.1)	
High (>32 times/month)	666 (36.7)	93 (22.9)	2 (0.6)	< 0.001
Milk and dairy products				
Low (<4 times/month)	594 (32.7)	188 (46.2)	67 (18.5)	
Moderate (4–12 times/month)	561 (30.9)	134 (32.9)	161 (44.4)	
High (>12 times/month)	659 (36.3)	85 (20.9)	135 (37.2)	< 0.001
Clinical (means±SDs)				
PV				
50-60 years (n=387)	27.3±13.4	30.0±13.7	23.1±8.2	0.007
60-70 years (n=977)	32.6±16.9	38.8±19.7	27.6±14.5	< 0.001
70–80 years (n=931)	35.4±21.4	42.2±21.5	34.5±27.5	0.004
≥80 years (n=289)	35.8±20.7	56.9±23.4	37.3±19.1	< 0.001
All (n=2584)	33.1±19.0	39.6±20.8	28.8±19.2	<0.001
IPSS				
50-60 years (n=387)	14.89±5.91	12.68±4.90	14.92±44.55	0.002
60-70 years (n=977)	15.96±6.10	13.86±5.43	17.90±5.57	<0.001
70–80 years (n=931)	16.80±6.59	15.95±6.12	21.67±5.46	<0.001
≥80 years (n=289)	18.26±6.39	19.32±6.36	22.55±8.05	0.066
All (n=2584)	16.44±6.37	14.84±5.95	18.65±5.96	< 0.001
IIEF-5				
50-60 years (n=387)	15 (8, 20)	19 (15, 21)	13 (11, 15)	<0.001
60–70 years (n=977)	12 (5, 17)	13 (9, 17)	9 (7, 11)	<0.001
70–80 years (n=931)	5 (0, 13)	3 (0, 8)	1 (0, 5)	< 0.001
≥80 years (n=289)	2 (0, 12)	1 (0, 5)	0 (0, 1)	0.003
All (n=2584)	9 (1, 15)	10 (3, 16)	8 (5, 12)	<0.001

Means and SDs were used to present normally distributed continuous variables, and medians (q1, q3) were used to present non-normally distributed variables (IIEF-5 score).

A p<0.05 by the Kruskal-Wallis test or one-way ANOVA was considered significant.

ANOVA, analysis of variance; BPH, benign prostatic hyperplasia; IIEF-5, International Index of Erectile Function 5; IPSS, International Prostate Symptom Score; LUTS, lower urinary tract symptoms; PV, prostate volume.

China and smallest among men residing in northwestern China. Comparisons between any two of the three regions were statistically significant (p<0.001) for all men and men 60–70 years of age. In contrast, IPSS showed an inverse distribution in the three geographic regions. It was highest in northwestern China and lowest in southern China. Comparisons between any two of the three geographic regions were nearly statistically significant with the exception of participants >80 years of age. Participants residing in northern and southern China had higher IIEF-5 scores than those residing in northwestern China (p<0.001 and p<0.01, respectively).

Tables 3 and 4 present the univariable and multivariable analyses for the PV, IPSS and IIEF-5 score. Considering the strong differences in diet across regions, the two factors were separated in two multivariate analysis models. First, based on the multivariable

linear analysis, PV enlarged as the age, BMI and vegetable intake increased ( $\beta$ =0.44, p<0.001;  $\beta$ =0.42, p<0.001;  $\beta$ =0.80, p<0.001). The association between PV and meat, fish and eggs intake failed to reach a statistical significance ( $\beta$ =0.05, p=0.711). Second, age ( $\beta$ =0.13, p<0.001), BMI ( $\beta$ =0.08, p<0.05) and cigarette smoking ( $\beta$ =0.89, p=0.001) independently increased IPSS. A higher level of education ( $\beta$ =-1.50, p<0.001) and more frequent meat, fish and eggs intake ( $\beta$ =-0.11, p<0.05) decreased IPSS. Third, age ( $\beta$ =-0.03, p<0.001), BMI ( $\beta$ =-0.02, p<0.001), low education level ( $\beta$ =0.10, p=0.025), vegetable intake ( $\beta$ =-0.04, p<0.001), and milk and dairy product intake ( $\beta$ =-0.08, p=0.001) decreased the IIEF-5 score. Meanwhile, regional differences were independently correlated with the PV, IPSS and IIEF-5 score, with the exception of participants in northern and southern China, who had similar IIEF-5 scores.

	PV		IPSS		IIEF-5	
Variables	Means±SDs	p Value	Means±SDs	p Value	Medians (q1, q3)	p Value
Education (years)						
≤9	35.3±20.6		17.84±6.50		8 (1, 14)	
>9	32.9±19.1	0.007	16.05±6.21	< 0.001	10 (2, 15)	0.001
Smoking						
No	33.2±19.4		16.14±6.02		8 (1, 14)	
Yes	34.1±19.7	0.250	17.04±6.49	< 0.001	10 (2, 15)	0.070
Alcohol consumption						
No	33.4±19.3		16.32±6.21		9 (1, 15)	
Yes	33.8±19.9	0.600	16.80±6.54	0.066	10 (2, 16)	0.002
Region						
Northern China	33.1±19.0		16.44±6.37		9 (1, 15)	
Southern China	39.6±20.8		14.84±5.95		10 (3, 16)	
Northwestern China	28.8±19.2	<0.001	18.65±5.96	< 0.001	8 (5, 12)	< 0.001
	β (95% CI)	р	β (95% CI)	р	β (95% CI)	р
Age (years)	0.44 (0.35 to 0.52)	<0.001	0.14 (0.11 to 0.16)	< 0.001	-0.35 (-0.38 to -0.32)	< 0.001
BMI (kg/m <sup>2</sup> )	0.48 (0.24 to 0.71)	<0.001	0.10 (0.03 to 0.18)	0.008	-0.20 (-0.28 to -0.12)	< 0.001
Meat, fish and eggs (times/week)	-0.01 (-0.25 to 0.26)	0.961	-0.13 (-0.21 to -0.05)	0.002	0.01 (-0.01 to 0.02)	0.269
Vegetables (times/week)	0.82 (0.47 to 1.16)	<0.001	0.09 (-0.02 to 0.21)	0.098	-0.04 (-0.06 to -0.02)	< 0.001
Milk and dairy products (times/week)	-0.02 (-0.10 to 0.14)	0.743	0.05 (0.01 to 0.09)	0.010	-0.01 (-0.20 to -0.01)	< 0.001

Means and SDs were used to present normally distributed continuous variables, and medians (q1, q3) were used to present non-normally distributed variables (IIEF-5 score). Categorical variables (education, smoking, alcohol consumption and region): t-test or one-way ANOVA was used for PV and IPSS; Wilcoxon test or Kruskal-Wallis test was used for the IIEF-5

Continuous variables (age, BMI and diet): univariate linear regressions was used for PV, IPSS and the IIEF-5 score.  $\beta$ : Unstandardised coefficients.

ANOVA, analysis of variance; BMI, body mass index; IIEF-5, International Index of Erectile Function 5; IPSS, International Prostate Symptom Score; PV, prostate volume.

Variables         β (95% CI)         p Value         β (95% CI)         p Value         β (95% CI)         p Value           Model 1 (included age, BMI, education, smoking, alcohol consumption and diet (weekly))         0.44 (0.35 to 0.53)         <0.001         0.03 (0.10 to 0.16)         <0.001         −0.03 (-0.04 to −0.02)         <0.001           Age (years)         0.44 (0.35 to 0.53)         <0.001         0.08 (0.01 to 0.16)         0.030         −0.02 (-0.04 to −0.01)         <0.001           BMI (kg/m²)         −1.60 (−3.40 to 0.20)         0.081         −1.50 (−2.08 to −0.91)         0.030         −0.02 (−0.04 to −0.01)         0.025           Smoking (no/yes)         −1.60 (−3.40 to 0.20)         0.081         −1.50 (−2.08 to −0.91)         0.001 (−0.01 to 0.18)         0.025           Alcohol consumption (no/yes)         0.72 (−0.96 to 2.39)         0.400         0.32 (−0.22 to 0.86)         0.246         0.01 (−0.07 to 0.09)         0.794           Meat, fish and eggs (times/week)         0.05 (−0.21 to 0.31)         0.711 (−0.18 to −0.01)         0.031 (−0.02 to 0.02)         0.01 (−0.02 to 0.02)         0.094           Vegetables (times/week)         0.00 (-0.21 to 0.31)         0.07 (−0.19 to 0.05)         0.272         0.06 (0.02 to 0.10)         0.074 (−0.02 to 0.02)         0.001 (−0.02 to 0.02)           Model La (adjusted for age, BMI, education, smoking,		PV		IPSS		IIEF-5 score	
diet (weekly))  <0.001  0.03 (0.01 to 0.16)  0.08 (0.01 to 0.16)  0.08 (0.01 to 0.16)  0.08 (0.01 to 0.16)  0.08 (0.01 to 0.16)  0.09 (0.01 to 0.16)  0.09 (0.02 to 0.09)  0.001  0.01 (0.001 to 0.01)  0.001  0.001 (0.001 to 0.01)  0.001 (0.001 to 0.01)  0.001 (0.001 to 0.01)  0.001 (0.001 to 0.01)  0.002 (0.001 to 0.01)  0.005 (0.001 to 0.02)  0.006 (0.02 to 0.10)  0.005 (0.001 to 0.02)  0.006 (0.02 to 0.10)  0.005 (0.014 to 0.05)  0.008 (0.014 to 0.05)	/ariables	β (95% CI)	p Value	β (95% CI)	p Value	β (95% CI)	p Value
<ul> <li>&lt;0.001 <li>0.13 (0.10 to 0.16)     <li>&lt;0.001 <li>0.08 (0.01 to 0.16)     <li>&lt;0.001 <li>0.08 (0.01 to 0.16)     </li> <li>0.081     </li> <li>0.081     </li> <li>-1.50 (-2.08 to -0.91)     </li> <li>0.081     </li> <li>0.89 (0.36 to 1.42)     </li> <li>0.001     </li> <li>0.001 (-0.04 to -0.01)     </li> <li>0.001 (-0.04 to 0.01)     </li> <li>0.001 (-0.01 to 0.18)</li> <li>0.001 (-0.01 to 0.08)</li> <li>0.001 (-0.08 to 0.08)</li> <li>0.001 (-0.07 to 0.09)</li> <li>0.001 (-0.07 to 0.09)</li> <li>0.001 (-0.07 to 0.09)</li> <li>0.002 (-0.02 to 0.20)</li> <li>0.127 (-0.04 (-0.05 to -0.02)</li> <li>0.005 (-0.02 to -0.02)</li> <li>0.005 (-0.04 to 0.02)</li> <li>0.005 (-0.13 to -0.04)</li> <li>0.006 (0.02 to 0.10)</li> <li>0.005 (-0.14 to 0.05)</li> <li>0.008 (-0.14 to 0.05)</li> <li>0.001 (-0.01 to 0.01)</li> <li>0.001 (-0.01 to 0.01)</li> </li></li></li></li></li></ul>	Model 1 (included age, BMI, education, sr	moking, alcohol consumption	and diet (weekl	y))			
<ul> <li>&lt;0.001 <li>0.08 (0.01 to 0.16)     <li>0.030</li> <li>0.02 (-0.04 to -0.01)</li> <li>0.081</li> <li>0.150 (-2.08 to -0.91)</li> <li>0.001</li> <li>0.10 (0.01 to 0.18)</li> <li>0.173</li> <li>0.89 (0.36 to 1.42)</li> <li>0.001</li> <li>0.001 (-0.08 to 0.08)</li> <li>0.246</li> <li>0.01 (-0.08 to 0.08)</li> <li>0.246</li> <li>0.01 (-0.07 to 0.09)</li> <li>0.711 (-0.18 to -0.01)</li> <li>0.031</li> <li>0.01 (-0.07 to 0.09)</li> <li>0.09 (-0.02 to 0.20)</li> <li>0.127</li> <li>0.004 (-0.05 to -0.02)</li> <li>0.005</li> <li>0.005</li> <li>0.005</li> <li>0.006 (0.02 to 0.10)</li> <li>0.005</li> <li>0.006 (-0.13 to -0.04)</li> <li>0.005</li> <li>0.014 to 0.05</li> <li>0.006 (-0.14 to 0.05)</li> <li>0.008</li> <li>2.49 (1.79 to 3.18)</li> <li>0.001</li> </li></li></ul>	Age (years)	0.44 (0.35 to 0.53)	<0.001	0.13 (0.10 to 0.16)	<0.001	-0.03 (-0.04 to -0.02)	<0.001
0.081	$BMI (kg/m^2)$	0.42 (0.19 to 0.65)	<0.001	0.08 (0.01 to 0.16)	0:030	-0.02 (-0.04 to -0.01)	<0.001
0.173	Education (≤9/>9 years)	-1.60 (-3.40 to 0.20)	0.081	-1.50 (-2.08 to -0.91)	<0.001	0.10 (0.01 to 0.18)	0.025
0.400 0.32 (-0.22 to 0.86) 0.246 0.01 (-0.07 to 0.09) 0.711 -0.11 (-0.18 to -0.01) 0.031 0.01 (-0.01 to 0.02) <0.001 0.09 (-0.02 to 0.20) 0.127 -0.04 (-0.05 to -0.02) 0.272 0.06 (0.02 to 0.10) 0.005 -0.08 (-0.13 to -0.04)  - ref  - ref - c0.001 -1.42 (-2.08 to -0.76) c0.001 -0.05 (-0.14 to 0.05) 0.008 2.49 (1.79 to 3.18) c0.001 -0.21 (-0.31 to -0.10)	Smoking (no/yes)	1.14 (-0.50 to 2.78)	0.173	0.89 (0.36 to 1.42)	0.001	-0.01 (-0.08 to 0.08)	0.979
0.711	Alcohol consumption (no/yes)	0.72 (-0.96 to 2.39)	0.400	0.32 (-0.22 to 0.86)	0.246	0.01 (-0.07 to 0.09)	0.794
<ul> <li><ul>              &lt;0.001</ul></li></ul>	Meat, fish and eggs (times/week)	0.05 (-0.21 to 0.31)	0.711	-0.11 (-0.18 to -0.01)	0.031	0.01 (-0.01 to 0.02)	0.095
0.272 0.06 (0.02 to 0.10) 0.005 -0.08 (-0.13 to -0.04)  - ref - ref - ref -0.08 to -0.76) <0.001 -0.05 (-0.14 to 0.05) <0.008 2.49 (1.79 to 3.18) <0.001 -0.21 (-0.31 to -0.10) <	Vegetables (times/week)	0.80 (0.46 to 1.14)	<0.001	0.09 (-0.02 to 0.20)	0.127	-0.04 (-0.05 to -0.02)	<0.001
- ref - ref - c0.001 -1.42 (-2.08 to -0.76) <0.001 -0.05 (-0.14 to 0.05) 0.008 2.49 (1.79 to 3.18) <0.001 -0.21 (-0.31 to -0.10) <	Milk and dairy products (times/week)	-0.07 (-0.19 to 0.05)	0.272	0.06 (0.02 to 0.10)	0.005	-0.08 (-0.13 to -0.04)	0.001
ref – ref – ref – ref 7.52 (5.48 to 9.57) <0.001 –1.42 (–2.08 to –0.76) <0.001 –0.05 (–0.14 to 0.05) nina –2.94 (–5.10 to –0.78) 0.008 2.49 (1.79 to 3.18) <0.001 –0.21 (–0.31 to –0.10) <	Model 2 (adjusted for age, BMI, education	i, smoking, alcohol consumption	ou)				
7.52 (5.48 to 9.57) <0.001	Northern China	ref	1	ref	1	ref	1
-2.94 (-5.10 to -0.78) 0.008 2.49 (1.79 to 3.18) <0.001 -0.21 (-0.31 to -0.10)	Southern China	7.52 (5.48 to 9.57)	<0.001	-1.42 (-2.08 to -0.76)	<0.001	-0.05 (-0.14 to 0.05)	0.369
	Northwestern China	-2.94 (-5.10 to -0.78)	0.008	2.49 (1.79 to 3.18)	<0.001	-0.21 (-0.31 to -0.10)	<0.001

Additionally, univariable and multivariable linear analyses of the PV, IPSS and IIEF-5 scores were performed in the 2584 participants with LUTS/BPH (see online supplementary table S1). A larger PV ( $\beta$ =-0.03, p<0.001) and a higher IPSS ( $\beta$ =-0.02, p=0.001) decreased the IIEF-5 score, and a larger PV resulted in a higher IPSS ( $\beta$ =0.04, p<0.001).

#### DISCUSSION

BMI, body mass index; IIEF-5, International Index of Erectile Function 5; IPSS, International Prostate Symptom Score; PV, prostate volume.

This study was the first with a relatively large sample size to investigate the effect of diet habits on LUTS and sexual function in Chinese men with LUTS/BPH from three different geographic regions. Our survey indicated that increased age, obesity and a low level of education are independently associated with an enlarged PV, more severe LUTS and worse male sexual dysfunction (MSD). Moreover, we also found that there were geographic variations with respect to PV, LUTS and MSD, which might partially be attributed to different dietary habits in different geographic regions.

Generally, the term clinical BPH is synonymous with LUTS in the absence of urethral strictures or prostate enlargement as detected on DRE. The LUTS metric most often used in epidemiological studies is the self-reported IPSS, which assesses self-reported severity of urinary symptoms and defines a moderate or severe condition as an IPSS $\geq$ 8. Also, the adverse effects of BPH are relatively apparent in men >50 years of age. According to the results of this study, among Chinese adults with LUTS/BPH, PV and LUTS deteriorated as age increased. Indeed, Shi *et al* and Chokkalingam *et al* selected age  $\geq$ 50 years as an inclusion criterion for studying BPH. Therefore, we selected LUTS/BPH cases among participants who were  $\geq$ 50 years of age and had an IPSS $\geq$ 8, and excluded non-BPH-associated LUTS cases.

The reported prevalence of LUTS/BPH varies widely in different studies due to the use of different measuring instruments, the heterogeneity of different study populations and population sampling methods. Chokkalingam et al's population-based study on West Africans reported the prevalence of DRE-detected prostate enlargement to be 62.3%. In total, 22.9% of 106 435 men >45 years of age had moderate-to-severe LUTS in Smith et al's<sup>20</sup> population-based study. Also, a prevalence of 38.2% for moderate-to-severe LUTS was reported among residents in China >65 years of age by Wong et al.2 In 2009, Xu et al21 reported a prevalence of 66.9% for BPH among those >50 years of age in Tianjin, China. Of note, the aforementioned studies all involved the general population and the age ranges of the participants were different. In this study, the general prevalence of LUTS/BPH was as high as 61.4% (2584/4208). The discrepancy between our results and others might be attributed to our selection of participants seeking a regular prostate check-up from the outpatient clinic. These participants might be more aware of LUTS or have LUTS due to BPH.

BMI is regarded as a health indicator, which would be influenced by dietary habits to some extent. Evidence supporting the influence of BMI on PV and LUTS was mixed. According to Schuurman et al,<sup>22</sup> an increase in BMI is associated with higher levels of oestrogen and oestradiol, and lower levels of plasma testosterone, which could deteriorate BPH or LUTS; however, according to Gupta et al<sup>13</sup> and Joseph et al,<sup>23</sup> no relationship between BMI and BPH or LUTS existed. Bhindi et al<sup>2</sup> reported that obesity is associated with increased PV, but not with worse LUTS. In this study, BMI was a risk factor for PV ( $\beta$ =0.42, p<0.001) and LUTS ( $\beta$ =0.08, p<0.05), with the exception of fluctuating IPSS in men with a BMI<18.5 kg/m<sup>2</sup>. The results of this study were consistent with those of Smith et al<sup>20</sup> and Hammersten and Hogstedt;<sup>14</sup> specifically, an increase in BMI is a risk factor for BPH or LUTS. In addition, we showed that MSD had a deteriorating trend as the BMI increased from  $18.5 \text{ kg/m}^2$  and at a low level ( $<18.5 \text{ kg/m}^2$ ), which was consistent with the results of Teles et  $al^{10}$  and Ghalayini et al, 11 who reported that the prevalence of ED increases in men who are overweight.

Interestingly, Zhao et al<sup>25</sup> also reported that elderly men of southern China (Guangdong) consumed relatively more meat food and less green vegetables than some northern cities (eg, Shandong), which might be attributed to the different dwellers' taste, cuisine areas and economy level in China. One of the clinically important findings of this study was the effect of diet habits on PV and LUTS, under the condition that a larger PV resulted in a higher IPSS (β=0.04, p<0.001) in the 2584 participants with LUTS/BPH (see online supplementary table S1). In other words, the association between the frequency of meat, fish and egg consumption and PV was not statistically significant (p=0.71); the frequency of meat, fish and egg consumption was inversely correlated to IPSS ( $\beta$ =-0.11, p<0.05). First, the reported influence of diet on PV is inconsistent. According to Zhang et al,26 the frequency of animal protein intake is positively correlated with PV. However, according to Shirazi et al, 19 no relationship was detected between dietary patterns and PV. Second, the relationship between diet and BPH is mixed. Bravi et al<sup>27</sup> and Suzuki et al<sup>28</sup> reported that men with high protein and low vegetable intake are at greater risk for developing BPH; however, self-reported measuring instruments were used to judge the status of BPH, and no information involving PV was available. In contrast, the results of Bhindi et  $al^{24}$  indicated that obesity, which is caused by a high-energy diet, is associated with a larger PV, but not worse LUTS. Moreover, Maserejian et al<sup>29</sup> claimed that the inverse association between protein intake and LUTS is supported by evidence that high-protein diets suppress sympathetic nervous system activity. The hormonal pathway, in which dietary protein can lower the plasma testosterone concentration or affect the oestradiol level, might also be a conceivable mechanism. Third, with respect to vegetables, we only showed that

the frequency of vegetable intake was positively associated with PV ( $\beta$ =0.80, p<0.001). Although most evidence supports the notion that increased vegetable intake can reduce the risk of BPH, no association existed between the frequency of vegetable intake and the risk for BPH according to Lagiou *et al.*<sup>16</sup> Also, we suspect that the difference in the proportion of vegetables consumption between Chinese and Western populations might have influenced the effect of vegetables on LUTS/BPH. The result that the frequency of milk and dairy products was positively associated with IPSS ( $\beta$ =0.06, p<0.05) was consistent with the previous studies. Therefore, uniform standards and more large-scale studies are needed to clarify the relationship between dietary habits and LUTS/BPH.

Additionally, Oksuz and Malhan<sup>30</sup> reported that men without diet-based lifestyle changes are at higher risk for MSD, and Khoo *et al*<sup>15</sup> found that a reduced-fat diet in obese Chinese men enhances male sexual function. In this study, however, vegetables and milk and dairy products were risk factors for MSD. These findings indicate that a balanced diet might benefit male sexual function. Of note, age ( $\beta$ =-0.03, p<0.001), PV ( $\beta$ =-0.03, p<0.001) and LUTS ( $\beta$ =-0.02, p=0.001) were all independent risk factors for MSD, which is consistent with previous studies.<sup>6-8</sup> <sup>12</sup> Thus, participants in our research were at high risk for MSD, which might explain the results of our study. More attention by clinicians should be paid to these groups of men to alleviate MSD.

Another clinically meaningful finding of our research was the dissociative trend of PV and LUTS in terms of geographic variation. In 1997, Yu et  $al^{31}$  conducted the survey on prevalence of prostatic hyperplasia in China, mainly relying on PV from DRE. According to the results of Yu et als<sup>32</sup> study, the population in south China had more severe prostate enlargement than that in northwest China, which is consistent with the results of our study. Although PV was positively associated with LUTS for the 2584 participants with LUTS/BPH (see online supplementary table S1), men with LUTS/BPH in south and northwest China had the mildest and most severe LUTS, respectively. First, regional differences in demographic characteristics like diet habits, education level, economic development, natural environment, etc, might contribute to the geographic variation of PV and LUTS to some extent.<sup>2</sup> <sup>3</sup> Second, morphological differences of prostate due to genetics and races might also act on the geographic distribution of PV and LUTS.<sup>32</sup> For instance, compared with the Caucasian population, Korean men have an incidence of LUTS similar to that in men in the Western hemisphere, despite having a smaller average PV. Third, BPH, especially in the early stage, might be more closely connected with voiding symptoms of LUTS. 17 34 Meanwhile, MSD had a similar geographical distribution as LUTS. Therefore, the assignment of public health policies for the authorities, as well as future research on LUTS/BPH and MSD for the epidemiologists, should be further performed.

Limitations affecting our current findings must be considered. First, sample size diversity existed in the three geographic regions. Meanwhile, our participants were native males ≥50 years of age who attended the clinic for a prostate check-up, who represented a more highly educated, higher health literate and higher socioeconomic group than the general population. Therefore, more community-based studies are imperative to reduce the selection bias. Second, the food frequency in our study, which classified the daily foods into three main categories, was designed to study the broad influence of the categories on PV, LUTS and MSD; we did not evaluate the associations between one single food and PV, LUTS and MSD.

#### CONCLUSION

The results of this study suggest that, in addition to factors including age, obesity and level of education, dietary habits and geographic differences might also play an important role in the variation of PV, LUTS and MSD for Chinese men with LUTS/BPH. Superficially, PV and LUTS might present a dissociative trend under the influence of diet habits and geographic variation.

#### **Author affiliations**

- <sup>1</sup>Department of Urology, Peking University First Hospital, Institute of Urology, Peking University, National Urological Cancer Center, Beijing, China
- <sup>2</sup>Department of Urology, Beijing Hospital, Beijing, China
- <sup>3</sup>Department of Urology, Beijing Friendship Hospital Affiliated with Capital Medical University, Beijing, China
- <sup>4</sup>Department of Urology, The First Affiliated Hospital of Xi'an Jiaotong University, Xi'an, China
- <sup>5</sup>Department of Urology, The Second Affiliated Hospital of Tianjin Medical University, Tianjin, China
- <sup>6</sup>Department of Urology, Sun Yat-Sen Memorial Hospital of Sun Yat-Sen University, Guangzhou, China
- <sup>7</sup>Department of Urology, Peking University People's Hospital, Beijing, People's Republic of China
- <sup>8</sup>Department of Urology, The Third Affiliated Hospital of Sun Yat-Sen University, Guangzhou, China
- <sup>9</sup>Department of Urology, Peking Union Medical College Hospital, Chinese Academy of Medical Sciences, Beijing, China
- <sup>10</sup>Department of Urology, Third Hospital of Peking University, Beijing, China
- <sup>11</sup>Department of Urology, Beijing Chao-yang Hospital Affiliated to Capital Medical University, Beijing, China
- <sup>12</sup>Department of Urology and Central Research Laboratory, Shandong University Second Hospital, Jinan, China
- <sup>13</sup>Minimally Invasive Urology Center, Provincial Hospital Affiliated to Shandong University, Jinan, China

**Acknowledgements** The authors thank Bang Zheng from School of Public Health, Peking University Health Science Center for his assistance with statistics analysis.

Contributors YC and WY are co-first authors who contributed equally to drafting and revising this manuscript. LZ and SW are the corresponding authors who contributed to designing the whole manuscript. YY, JW, YT, DH, YX, JH, XW, XG, HL, LM, NZ, SZ and XJ contributed equally to collecting and sorting the data.

**Funding** This work was supported by Grants from the "The Research Special Fund for Public Welfare Industry of Health" (grant number: 201002010).

Competing interests None declared.

#### Patient consent Obtained.

Ethics approval The study was approved by the Institutional Review Board of The First, Second, and Third Affiliated Hospitals of Peking University, Peking Union Medical College Hospital, Beijing Chao-Yang Hospital, Beijing Friendship Hospital, Beijing Hospital, The Second Affiliated Hospital of Tianjin Medical University, The Second and Third Affiliated Hospitals of Zhongshan University, and The First Affiliated Hospitals of Xi'an Jiaotong University.

Provenance and peer review Not commissioned; externally peer reviewed.

Data sharing statement No additional data are available.

Open Access This is an Open Access article distributed in accordance with the Creative Commons Attribution Non Commercial (CC BY-NC 4.0) license, which permits others to distribute, remix, adapt, build upon this work non-commercially, and license their derivative works on different terms, provided the original work is properly cited and the use is non-commercial. See: http://creativecommons.org/licenses/by-nc/4.0/

#### **REFERENCES**

- Tewari R, Rajender S, Natu SM, et al. Diet, obesity, and prostate health: are we missing the link? J Androl 2012;33:763–76.
- Wong SY, Woo J, Hong A, et al. Risk factors for lower urinary tract symptoms in southern Chinese men. *Urology* 2006;68:1009–14.
- Eckhardt MD, van Venrooij GE, Boon TA. Symptoms, prostate volume, and urodynamic findings in elderly male volunteers without and with LUTS and in patients with LUTS suggestive of benign prostatic hyperplasia. *Urology* 2001;58:966–71.
   Yang HJ, Doo SW, Yang WJ, *et al.* Which obesity index best
- Yang HJ, Doo SW, Yang WJ, et al. Which obesity index best correlates with prostate volume, prostate-specific antigen, and lower urinary tract symptoms? *Urology* 2012;80:187–90.
- Zucchetto A, Tavani A, Dal Maso L, et al. History of weight and obesity through life and risk of benign prostatic hyperplasia. Int J Obes (Lond) 2005;29:798–803.
- Martin SA, Atlantis E, Lange K, et al. Predictors of sexual dysfunction incidence and remission in men. J Sex Med 2014;11:1136–47.
- Braun M, Wassmer G, Klotz T, et al. Epidemiology of erectile dysfunction: results of the "Cologne Male Survey". Int J Impot Res 2000;12:305–11.
- Berrada S, Kadri N, Mechakra-Tahiri S, et al. Prevalence of erectile dysfunction and its correlates: a population-based study in Morocco. Int J Impot Res 2003;15:3–7.
- Wang GC, Zheng JH, Yang B, et al. Impacts of histological prostatitis on sexual function and lower urinary tract symptoms in patients with benign prostatic hyperplasia. *Urology* 2013;82:1094–7.
- Teles AG, Carreira M, Alarcão V, et al. Prevalence, severity, and risk factors for erectile dysfunction in a representative sample of 3,548 Portuguese men aged 40 to 69 years attending primary healthcare centers: results of the Portuguese erectile dysfunction study. J Sex Med 2008;5:1317–24.
- Ghalayini IF, Al-Ghazo MA, Al-Azab R, et al. Erectile dysfunction in a Mediterranean country: results of an epidemiological survey of a representative sample of men. Int J Impot Res 2010;22:196–203.
- Fwu CW, Kirkali Z, McVary KT, et al. Cross-sectional and longitudinal associations of sexual function with lower urinary tract symptoms in men with benign prostatic hyperplasia. J Urol 2015;193:231–8.
- Gupta A, Gupta S, Pavuk M, et al. Anthropometric and metabolic factors and risk of benign prostatic hyperplasia: a prospective cohort study of Air Force veterans. *Urology* 2006;68:1198–205.
- Hammersten J, Hogstedt B. Hyperinsulinaemia as a risk factor for developing benign prostatic hyperplasia. Eur Urol 2001;39:151–8.
- Khoo J, Ling PS, Tan J, et al. Comparing the effects of meal replacements with reduced-fat diet on weight, sexual and endothelial function, testosterone and quality of life in obese Asian men. Int J Impot Res 2014;26:61–6.
- Lagiou P, Wuu J, Trichopoulou A, et al. Diet and benign prostatic hyperplasia: a study in Greece. *Urology* 1999;54:284–90.
  Chokkalingam AP, Yeboah ED, Demarzo A, et al. Prevalence of
- Chokkalingam AP, Yeboah ED, Demarzo A, et al. Prevalence of BPH and lower urinary tract symptoms in West Africans. Prostate Cancer Prostatic Dis 2012;15:170–6.
- Shi J, Sun Z, Cai T, et al. Development and validation of a quality-of-life scale for Chinese patients with benign prostatic hyperplasia. BJU Int 2004;94:837–44.
- Shirazi M, Ariafar A, Zeyghami S, et al. Association of diet with prostate specific antigen and prostate volume. Nephrourol Mon 2014;6:e19411.
- Smith DP, Weber MF, Soga K, et al. Relationship between lifestyle and health factors and severe lower urinary tract symptoms (LUTS)

- in 106,435 middle-aged and older Australian men: population-based study. *PLoS ONE* 2014;9:e109278.
- Xu Y, Zhang ZH, Cheng R, et al. Prevalence of benign prostate hyperplasia and its relative factors in rural areas of Tianjin in 2008. Chin J Urol 2009;30:761–4.
- Schuurman AG, Goldbohm RA, Dorant E, et al. Anthropometry in relation to prostate cancer risk in the Netherlands Cohort Study. Am J Epidemiol 2000;151:541–9.
- Joseph MA, Harlow SD, Wei JT, et al. Risk factors for lower urinary tract symptoms in a population-based sample of African-American men. Am J Epidemiol 2003;157:906–14.
- Bhindi B, Margel D, Trottier G, et al. Obesity is associated with larger prostate volume but not with worse urinary symptoms: analysis of a large multiethnic cohort. *Urology* 2014;83:81–7.
- Zhao W, You Y, Zhang X, et al. Study on the food consumption pattern of elderly men in four "cuisine" areas of China. Wei Sheng Yan Jiu 2002;31:34–7.
- Zhang SX, Yu B, Guo SL, et al. Comparison of incidence of BPH and related factors between urban and rural inhabitants in district of Wannan. Zhonghua Nan Ke Xue 2003;9:45–7.
- Bravi F, Bosetti C, Dal Maso L, et al. Food groups and risk of benign prostatic hyperplasia. *Urology* 2006;67:73–9.

- Suzuki S, Platz EA, Kawachi I, et al. Intakes of energy and macronutrients and the risk of benign prostatic hyperplasia. Am J Clin Nutr 2002;75:689–97.
- Maserejian NN, Giovannucci EL, McKinlay JB. Dietary macronutrients, cholesterol, and sodium and lower urinary tract symptoms in men. *Eur Urol* 2009;55:1179–89.
- Oksuz E, Malhan S. The prevalence of male sexual dysfunction and potential risk factors in Turkish men: a web-based survey. Int J Impot Res 2005;17:539

  –45.
- 31. Yu P, Zheng H, Su H, *et al.* Prevalence of prostatic hyperplasia and its relative factors in six cities of China in 1997. *Zhonghua Liu Xing Bing Xue Za Zhi* 2000;21:276–9.
- Yu EY, Siegal JA, Meyer GE, et al. Histologic differences in benign prostate hyperplasia between Chinese and American men. Prostate 1997;31:175–9.
- Lee É, Yoo KY, Kim Y, et al. Prevalence of lower urinary tract symptoms in Korean men in a community-based study. Eur Urol 1998;33:17–21.
- Lee SH, Chung BH, Kim CS, et al. Survey on benign prostatic hyperplasia distribution and treatment patterns for men with lower urinary tract symptoms visiting urologists at general hospitals in Korea: a prospective, noncontrolled, observational cohort study. Urology 2012;79:1379–84.