Dietary intake habits and the prevalence of nocturia in Japanese patients with type 2 diabetes mellitus

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Keywords

Dietary behavior, Nocturia, Vegetable

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ABSTRACT

Aims/Introduction: No reports have been published on the association between dietary intake habits and nocturia in the diabetes population. We therefore evaluated this issue among Japanese patients with diabetes mellitus.

Materials and Methods: Study participants in the present study were 785 Japanese patients with type 2 diabetes mellitus. Self-administered questionnaires were used to assess each type of dietary intake habit. Vegetable intake habit was assessed by the following question: "Do you have vegetables or seaweed every day?" We used the following two outcomes: (i) nocturia: >2 voids per night; and (ii) severe nocturia: >3 voids per night. Adjustment was made for age, sex, body mass index, glycated hemoglobin, hypertension, dyslipidemia, smoking, drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy.

Results: The prevalence of nocturia, severe nocturia, and vegetable intake habit was 39.9%, 14.4% and 67.3%, respectively. After adjusting for confounding factors, vegetable intake habit was independently inversely associated with nocturia and severe nocturia: the adjusted odds ratios were 0.67 (95% confidence interval [CI] 0.48-0.94) and 0.46 (95% CI 0.30-0.71), respectively. Among male patients, vegetable intake habit was independently inversely associated with severe nocturia, but not nocturia: the adjusted OR was 0.51 (95% Cl 0.29–0.88). Among female patients, vegetable intake habit was independently inversely associated with nocturia and severe nocturia: the adjusted ORs were 0.44 (95% Cl 0.24-0.79) and 0.34 (95% CI 0.15-0.78), respectively.

Conclusions: We found an inverse association between vegetable intake habit and nocturia in Japanese patients with type 2 diabetes mellitus.

INTRODUCTION

Nocturia, one type of voiding dysfunction, negatively affects nocturnal sleep, daytime sleepiness and quality of life. Nocturia is common among patients with diabetes mellitus. Among Japanese patients with type 2 diabetes mellitus, nocturia was

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significantly positively associated with erectile dysfunction¹, diabetic retinopathy² and depressive symptoms³.

Among the general population, some evidence exists regarding the relationship between dietary intake and voiding dysfunction including lower urinary tract symptoms (LUTS), overactive bladder (OAB) and benign prostate hyperplasia (BPH). Intake levels of vegetables⁴⁻⁹, vegetable fat⁶,

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isoflavones⁷ and citrus juice⁸ were significantly inversely associated with LUTS, whereas a significant positive relationship between total energy intake and LUTS was found⁹. Inverse associations between the intake of vegetables, bread and chicken and OAB were observed¹⁰. Intake levels of vitamin D, protein and potassium, as well as beer intake, were significantly inversely associated with the onset of OAB^{11,12}. Intake levels of vegetables and vegetable fat were significantly inversely associated with BPH^{7,13–16}.

No data regarding the association between dietary intake and nocturia are available among patients with type 2 diabetes mellitus. The importance of research into the possible role of dietary factors in voiding dysfunction is emphasized by the fact that diet is modifiable. The aim of the present study was to evaluate the association between dietary intake habits and nocturia among Japanese patients with type 2 diabetes mellitus.

METHODS

Study population

The Dogo Study is an ongoing multicenter prospective cohort study of patients with diabetes. All 1051 Japanese patients with type 2 diabetes mellitus who gave informed consent were enrolled from the participating local base hospitals in Ehime prefecture. The patients in this cohort study were recruited from 2009 to 2014. The final analysis sample in this study consisted of 785 patients, as 266 patients were excluded due to incomplete data. The present study protocol conforms to the ethical guidelines of the 1975 Declaration of Helsinki as reflected in a priori approval by the institutional review board of the Ehime University Graduate School of Medicine.

Measurements

Each participant completed a self-administered questionnaire, which collected data on the variables. Information on medication was based on admission records, medical records and/or a questionnaire. Exercise habit was defined as positive if the study participants reported exercising 60 min or more per week during leisure time, regardless of the exercise intensity. The definitions of variables were previously described in detail^{1–3}. The definitions of microvascular diseases were based on the Classification of Diabetic Nephropathy 2014^{17,18}, the Fukuda standard¹⁹ and the abbreviated diagnosis of diabetic neuropathy²⁰.

Assessment of dietary intake habits

We used a self-administered questionnaire to estimate dietary intake using the following questions: (i) "Do you have fried food every day?"; (ii) "Do you have one or more eggs every day?"; (iii) "Do you have meat more than twice a week?"; (iv) "Do you have fish or shellfish more than twice a week?"; (v) "Do you have salty pickles more than twice a day?"; (vi) "Do you have vegetables or seaweed every day?"; (vii) "Do you have fruit every day?"; (viii) "Do you have soy products every day?"; and (ix) "Do you consume dairy products every day?". Patients answered either "yes" or "no"^{21,22}.

Assessment of nocturia

The definition of nocturia was based on a questionnaire regarding the numbers of voids^{1–3}. We used the following outcomes: (i) nocturia was defined as ≥ 2 voids per night; and (ii) severe nocturia was defined as ≥ 3 voids per night.

Statistical analysis

We selected age, sex, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current smoking, current drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy as a priori as potential confounding factors. Logistic regression analyses were carried out to estimate crude odds ratios (ORs) and their 95% confidence intervals (CIs) for nocturia according to dietary intake habits. Additionally, multiple logistic regression analysis was carried out to adjust for confounding factors. All computations were carried out using SAS software package version 9.4 (SAS Institute Inc., Cary, NC, USA).

Table 1 | Clinical characteristics of the 785 study participants

Variable	n (%)
Age, years (mean ± SD)	61.7 ± 11.2
Male (%)	489 (62.3)
BMI, kg/m ² (mean ± SD)	25.2 ± 4.8
HbA1c, % (mean ± SD)	7.89 ± 1.83
Duration of type 2 diabetes mellitus, years (mean \pm SD)	11.0 ± 10.2
Current drinking (%)	323 (41.2)
Current smoking (%)	144 (18.3)
Exercise habit	258 (32.9)
Use of diuretics	89 (11.3)
Hypertension (%)	551 (70.2)
Dyslipidemia (%)	585 (74.5)
Diabetic neuropathy (%)	483 (61.5)
Diabetic retinopathy (%)	231 (29.4)
Diabetic nephropathy (%)	80 (10.2)
Stroke (%)	53 (6.8)
Ischemic heart disease (%)	89 (11.3)
Nocturia	
Nocturia, ≥2 voids per night (%)	313 (39.9)
Severe nocturia, ≥3 voids per night (%)	113 (14.4)
Dietary intake habits	
Fried foods (%)	174 (24.7)
Eggs (%)	329 (41.9)
Fatty meat (%)	238 (30.3)
Seafood (%)	553 (70.5)
Pickles (%)	248 (31.6)
Vegetables (%)	528 (67.3)
Fruit (%)	432 (55.0)
Soy products (%)	473 (60.3)
Dairy products (%)	460 (58.6)

BMI, body mass index; Hb1Ac, glycated hemoglobin; SD, standard deviation.

RESULTS

Characteristics of the 785 patients are listed in Table 1. The prevalence of nocturia and severe nocturia was 39.9% and 14.4%, respectively. Dietary intake habits of fried foods, eggs, fatty meat, seafood, pickles, vegetables, fruit, soy products, and dairy products were reported at 24.7%, 41.9%, 30.3%, 70.5%, 31.6%, 67.3%, 55.0%, 60.3% and 58.6% of study participants, respectively. The percentage intake of fish, fruit, soy products and dairy products among patients with vegetable intake habit was significantly low among all patients (data not shown). Table 2 shows the crude and adjusted ORs and 95% CIs for nocturia in relation to dietary intake habits among the 785 patients. In the crude analysis, seafood, fruit and dairy products intake habits were positively associated with nocturia. The association between seafood, fruit, and dairy products and nocturia completely disappeared after adjusting for age, sex, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current

 $\begin{array}{c|c} \textbf{Table 2} & \mbox{ Crude and adjusted odds ratios and 95\% confidence} \\ \mbox{ intervals for nocturia in relation to dietary intake habits among all } \\ \mbox{ patients} \end{array}$

smoking, current drinking, exercise habits, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy. After adjustment, vegetable intake habit was independently inversely associated with nocturia: the adjusted OR was 0.67 (95% CI 0.48-0.94). Table 3 shows the crude and adjusted ORs and 95% CIs for severe nocturia in relation to dietary intake habits. In crude analysis, fruit and daily products intake habits were positively associated with severe nocturia, whereas vegetable intake was inversely associated with severe nocturia. After adjustment, vegetable intake habit was independently inversely associated with severe nocturia: the adjusted OR was 0.46 (95% CI 0.30-0.71). Among male patients, vegetable intake habit was independently inversely associated with severe nocturia, but not nocturia: the adjusted OR was 0.51 (95% CI 0.29-0.88; Table 4). Among female patients, vegetable intake habit was independently inversely associated with nocturia and severe nocturia: the adjusted ORs

Table 3 Crude and adjusted odds ratios and 95% confidence
intervals for severe nocturia in relation to dietary intake habits among
all patients

Prevalence (%)

88/591 (14.9)

25/194 (12.9)

71/456 (15.6)

42/329 (12.8)

79/547 (14.4)

34/238 (14.3)

26/232 (11.2)

87/553 (15.7)

202/537 (37.6)

111/248 (44.8)

51/257 (19.8)

62/528 (11.7)

39/353 (11.1)

74/432 (17.1)

122/312 (39.1)

191/473 (40.4)

41/325 (12.6)

72/460 (15.7)

Variable

Severe nocturia Fried foods (%)

No

Yes Eggs (%) No

Yes

Yes

Yes

Yes

Fruit (%) No

Yes

Yes

No

Yes

Soy products (%) No 122/3

Dairy products (%)

Yes

Pickles (%) No

Vegetables (%) No 5

Seafood (%) No

Fatty meat (%) No 7 Crude OR

(95% CI)

100

100

1.00

1.00

1.00

1.00

1.00

1.00

1.00

0.85 (0.52-1.34)

0.79 (0.52-1.19)

0.99 (0.63-1.51)

1.48 (0.94-2.40)

1.22 (0.78-1.85)

0.54 (0.36-0.81)

1.66 (1.10-2.54)

1.00 (0.67-1.50)

1.29 (0.86-1.96)

Adjusted OR

0.99 (0.58-1.65)

0.87 (0.56-1.33)

1.16 (0.72–1.84)

1.09 (0.72-1.84)

1.05 (0.67-1.64)

0.46 (0.30-0.71)

1.31 (0.83-2.09)

0.99 (0.64-1.54)

1.11 (0.72–1.73)

(95% CI)

100

1.00

1.00

1.00

1.00

1.00

1.00

1.00

1.00

Variable	Prevalence (%)	Crude OR (95% CI)	Adjusted OR (95% Cl)
Nocturia			
Fried foc	ods (%)		
No	242/5,941 (41.0)	1.00	1.00
Yes	71/194 (36.6)		1.12 (0.77-1.61)
Eggs (%)		x 2	
No	185/456 (40.6)	1.00	1.00
Yes	128/329 (38.9)	0.93 (0.70–1.25)	1.03 (0.76-1.40)
Fatty me	at (%)		
No	217/547 (39.7)	1.00	1.00
Yes	96/238 (40.3)	1.03 (0.75–1.40)	1.33 (0.94–1.87)
Seafood	(%)		
No	74/232 (31.9)	1.00	1.00
Yes	239/553 (43.2)	1.63 (1.18–2.25)	1.22 (0.86–1.73)
Pickles (9	6)		
No	202/537 (37.6)	1.00	1.00
Yes	111/248 (44.8)	1.34 (0.99–1.82)	1.11 (0.79–1.54)
Vegetabl	es (%)		
No	110/257 (42.8)	1.00	1.00
Yes	203/528 (38.5)	0.84 (0.62–1.13)	0.67 (0.48-0.94)
Fruit (%)			
No	121/353 (34.3)	1.00	1.00
Yes	192/432 (44.4)	1.53 (1.15–2.05)	1.08 (0.78–1.50)
Soy proc	lucts (%)		
No	122/312 (39.1)	1.00	1.00
Yes	191/473 (40.4)	1.06 (0.79–1.41)	0.95 (0.69–1.30)
Dairy pro	oducts (%)		
No	113/325 (34.8)	1.00	1.00
Yes	200/460 (43.5)	1.44 (1.08–1.94)	1.25 (0.91–1.71)

Odds ratios (OR) were adjusted for age, sex, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current smoking, current drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy. Cl, confidence interval. Odds ratios (OR) were adjusted for age, sex, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current smoking, current drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy. Cl, confidence interval.

Table 4 Crude and adjusted odds ratios and 95% confidence
intervals for severe nocturia in relation to dietary intake among male
patients

Variable	Prevalence (%)	Crude OR (95% CI)	Adjusted OR (95% CI)
Nocturia			
Fried foc	ods (%)		
No	151/350 (43.1)	1.00	1.00
Yes	55/139 (39.6)	086 (0.58–1.29)	1.26 (0.80–1.99
Eggs (%)			
No	122/273 (44.7)	1.00	1.00
Yes	84/216 (38.9)	0.79 (0.55–1.13)	0.93 (0.63–1.38
Fatty me	eat (%)		
Ńo	143/344 (41.6)	1.00	1.00
Yes	63/145 (43.5)	1.08 (0.73-1.60)	1.46 (0.93-2.30
Seafood		× ,	x
No	49/145 (33.8)	1.00	1.00
Yes	157/344 (45.6)	1.65 (1.10-2.48)	1.16 (0.75–1.81
Pickles (9			
No	132/336 (39.3)	1.00	1.00
Yes	74/153 (48.4)	1.45 (0.98–2.13)	1.20 (0.79–1.83
Vegetabl			
No	74/178 (41.6)	1.00	1.00
Yes	132/311 (42.4)	1.04 (0.71–1.51)	0.81 (0.53–1.23
Fruit (%)	102,011 (1211)		0101 (0100 1120
No	90/241 (37.3)	1.00	1.00
Yes		1.47 (1.03–2.12)	0.93 (0.61–1.40
Soy proc		1.17 (1.03 2.12)	0.55 (0.01 1.10
No	88/212 (41.5)	1.00	1.00
Yes	118/277 (42.6)	1.05 (0.73–1.50)	0.91 (0.61–1.35
	oducts (%)	1.05 (0.75 1.50)	0.91 (0.01 1.95
No	82/219 (37.4)	1.00	1.00
Yes	124/270 (45.9)	1.42 (0.99–2.05)	1.12 (0.75–1.66
Severe noc		1.12 (0.99 2.03)	1.12 (0.75 1.00
Fried foc			
No	51/336 (15.2)	1.00	1.00
Yes	28/153 (18.3)	0.71 (0.39–1.22)	0.83 (0.43–1.55
Eggs (%)		0.71 (0.35 1.22)	0.00 (001 1.00
No	50/273 (18.3)	1.00	1.00
Yes	29/216 (13.4)	0.69 (0.42–1.13)	0.83 (0.48–1.40
Fatty me		0.09 (0.42-1.13)	0.00 (0.40-1.40
No	54/344 (15.7)	1.00	1.00
Yes	25/145 (17.2)	1.22 (0.66–1.86)	1.19 (0.65–2.14
Seafood		1.22 (0.00-1.00)	1.19 (0.00–2.14
No	17/145 (11.7)	1.00	1.00
Yes			1.14 (0.62–2.17
	62/344 (18.0)	1.66 (0.95–3.02)	1.14 (0.02–2.17
Pickles (%		1.00	1.00
No	51/336 (15.2)	1.00	1.00
Yes	28/153 (18.3)	1.25 (0.74–2.06)	1.15 (0.65–2.00
Vegetabl		1.00	1.00
No	36/178 (20.2)	1.00	1.00
Yes	43/311 (13.8)	0.63 (0.39–1.03)	0.51 (0.29–0.88
Fruit (%)	22/241 /12 2	1.00	1.00
No	32/241 (13.3)	1.00	1.00
Yes	47/248 (19.0)	1.53 (0.94–2.51)	0.98 (0.56–1.72

Table 4 (Continued)

Variable	Prevalence (%)	Crude OR (95% CI)	Adjusted OR (95% Cl)
Soy proc	lucts (%)		
No	32/212 (15.1)	1.00	1.00
Yes	47/277 (17.0)	1.15 (0.71–1.89)	1.12 (0.66–1.93)
Dairy pro	oducts (%)		
No	30/219 (13.7)	1.00	1.00
Yes	49/270 (18.2)	1.40 (0.86–2.31)	1.05 (0.62–1.81)

Odds ratios (OR) were adjusted for age, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current smoking, current drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy. Cl, confidence interval.

were 0.44 (95% CI 0.24–0.79) and 0.34 (95% CI 0.15–0.78), respectively (Table 5). In sensitive analysis, the use of diuretics was independently positively associated with nocturia, but not severe nocturia in all patients. Among patients with good glucose control (glycated hemoglobin <6.5%), the 1.5-anhydroglucitol level among patients with vegetable intake habit was significantly higher than that without.

DISCUSSION

To our knowledge, the present study is the first to show a significant inverse association between vegetable intake habit and nocturia and severe nocturia among Japanese patients with type 2 diabetes mellitus.

In the general population, some evidence of the association between intake of vegetables and voiding dysfunction exists. Vegetable intake was significantly inversely associated with LUTS in a Chinese study of 1,564 men⁴, in a Finnish study of 3,143 men⁵, in a Chinese study of 2,000 men⁷ and in three USA studies^{6,8,9}. Similarly, in a USA study of 7,043 women, vegetable intake was significantly inversely associated with OAB¹⁰. In USA studies of 51,529 men⁶, 4,770 men¹³, 32,265 men14 and in a case-control study of Japanese men15, an inverse association between vegetable intake and BPH was reported. In contrast, in a Greek study of 430 men, vegetable intake was not associated with BPH23. The present results are partially consistent with those of the previous studies regarding voiding dysfunction. The discrepancies among these studies might be explained, at least in part, by differences in sample size; the definition of voiding dysfunction; characteristics, such as age, sex, race, body mass index, prevalence of stroke and hypertension; and other confounding factors.

The associations between several types of dietary intake excluding vegetables and voiding dysfunction were observed. In a UK study of 7,043 women, intake of bread and chicken was significantly inversely associated with the onset of OAB¹⁰. In a USA study of 1,545 men aged 30–79 years, intake of total energy and sodium were significantly positively associated with

Table 5 Crude and adjusted odds ratios and 95% confidence
intervals for severe nocturia in relation to dietary intake among female
patients

Variable	Prevalence (%)	Crude OR (95% CI)	Adjusted OR (95% Cl)
Nocturia			
Fried foc	ods (%)		
No	91/241 (37.8)	1.00	1.00
Yes	16/55 (29.1)	0.68 (0.35-1.26)	0.86 (0.43-1.67)
Eggs (%)			
No	63/183 (34.4)	1.00	1.00
Yes	44/113 (38.9)	1.22 (0.75–1.97)	1.29 (0.77–2.17)
Fatty me			
No	74/203 (36.5)	1.00	1.00
Yes	33/93 (35.5)	0.96 (0.57–1.59)	1.13 (0.65–1.95)
Seafood			
	28/87 (28.7)	1.00	1.00
	82/209 (39.2)	1.60 (0.94–2.79)	1.31 (0.73–2.39)
Pickles (9			
No	70/201 (34.8)	1.00	1.00
Yes	37/95 (39.0)	1.19 (0.72–1.97)	0.97 (0.56–1.65)
Vegetabl			
	36/79 (45.6)	1.00	1.00
	71/217 (32.7)	0.58 (0.34–0.99)	0.44 (0.24–0.79)
Fruit (%)			
No	31/112 (27.7) 76/184 (41.3)	1.00	1.00
Yes	76/184 (41.3)	1.84 (1.11–3.08)	1.33 (0.75–2.35)
Soy proc		1.00	1.00
	34/100 (34.0)	1.00	1.00
	73/296 (37.2)	1.15 (0.70–1.92)	1.03 (0.60–1.79)
	oducts (%)	1.00	1.00
	31/106 (29.3)	1.00	1.00
	76/190 (40.0)	1.61 (0.98–2.71)	1.63 (0.96–2.82)
Severe noc			
Fried foc		1.00	1.00
	27/241 (11.2)	1.00	1.00
	7/55 (12.7)	1.16 (0.44–2.68)	1.35 (0.49–3.33)
Eggs (%)	21/183 (11.5)	1.00	1.00
NO	13/113 (11.5)	1.00	1.00
		1.00 (0.47–2.07)	1.00 (0.45–2.13)
Fatty me No	25/203 (12.3)	1.00	1.00
	25/205 (12.5) 9/93 (9.7)	0.76 (0.32–1.65)	1.00 0.77 (0.32–1.74)
Yes Seafood		0.70 (0.52-1.05)	0.77 (0.52–1.74)
No	9/87 (10.3)	1.00	1.00
Yes	25/209 (12.0)	1.18 (0.54–2.77)	0.98 (0.42-2.45)
Pickles (9		1.10 (0.54-2.77)	0.90 (0.42-2.43)
No	22/201 (11.0)	1.00	1.00
Yes	12/95 (12.6)	1.18 (0.54–2.45)	0.88 (0.39–1.92)
Vegetabl		1.10 (0.57-2.75)	0.00 (0.39-1.92,
No	15/79 (19.0)	1.00	1.00
Yes	19/217 (8.8)	0.41 (0.20–0.86)	0.34 (0.15–0.78)
Fruit (%)	12/21/ (0.0)	0.11 (0.20 0.00)	0.01 (0.10 0.70)
No	7/112 (6.3)	1.00	1.00
Yes	27/184 (14.7)	2.58 (1.14–6.63)	2.20 (0.88–6.11)
	277101(1177)	2.50 (1.11 0.05)	2,20 (0.00 0.11

Table 5 (Continued)

Variable	Prevalence (%)	Crude OR (95% CI)	Adjusted OR (95% Cl)
Soy prod	ucts (%)		
No	13/100 (13.0)	1.00	1.00
Yes	21/196 (10.7)	0.80 (0.39–1.72)	0.80 (0.37–1.80)
Dairy pro	ducts (%)		
No	11/106 (10.4)	1.00	1.00
Yes	23/190 (12.1)	1.19 (0.57–2.64)	1.13 (0.51–2.59)

Odds ratios (OR) were adjusted for age, body mass index, glycated hemoglobin, hypertension, dyslipidemia, current smoking, current drinking, exercise habit, stroke, ischemic artery disease, diabetic nephropathy, diabetic neuropathy and diabetic retinopathy. Cl, confidence interval.

LUTS⁹. Dietary isoflavone intake was significantly inversely associated with LUTS in a Chinese study of 2,000 men⁷. Intake of citrus juice was significantly inversely associated with LUTS in a USA study of 4,144 participants⁸. Beer intake was significantly inversely associated with the onset of OAB in a UK study of men¹².

No evidence regarding the association between dietary intake and nocturia exists. Nocturia is a storage symptom. In the general population, citrus juice was significantly inversely associated with storage symptoms in men, but not women; coffee intake was significantly positively associated with storage symptoms only in men, while soda intake was significantly positively associated with storage symptoms only in women⁸.

The mechanism linking vegetable intake habit and nocturia is not well understood. Vegetables are high in anti-oxidants, fibers, minerals and vitamins²⁴. Vegetable intake might reduce postprandial hyperglycemia²⁵. Among patients with diabetes, postprandial hyperglycemia is associated with increasing oxidative stress and inflammation^{26,27}. Vegetable intake might beneficially affect the prostate and bladder by inhibiting inflammation and oxidative damage.

The present study had several limitations that must be acknowledged. First, we used a cross-sectional analysis to assess the association between dietary intake habits and nocturia. Second, we used a "yes"/"no" response format partially validated questionnaire to assess dietary intake habits. A previous study using the same "yes"/"no" format questionnaire found that the participants who reported eating something frequently had a higher intake than the other participants²². In previous epidemiological studies, the participants with a high vegetable intake were older, and showed a lower prevalence of hypertension and current smoking^{28,29}. In the present study, similar characteristics as previous studies were observed among patients with vegetable intake habit. Therefore, we might assess the vegetable intake habit by this questionnaire. A "yes"/"no" response format questionnaire should be used only to assess vegetable intake habits. As consuming vegetables is thought to be a healthy habit, thus we should consider the possibility of misclassification. However, the possibility of non-differential outcome misclassification might have biased the magnitude of the observed associations toward the null. The questionnaire in the present study was inadequate to evaluate total energy intake, nutrient intake, and the dose-response association between eating vegetables and nocturia. Thus, further research based on a validated food frequency questionnaire is required to confirm the association between intake of vegetables and nocturia. Third, we used a questionnaire to survey nocturia. We could not carry out a urological test for nocturia, nor could we estimate BPH or bladder function. In a previous study, the overestimation of nocturia based on a questionnaire was reported³⁰. Among previous epidemiological studies⁴⁻¹⁶, however, the definition of nocturia was based on a questionnaire and/or interview. Fourth, many of the patients might have opted to consume vegetables to control their hyperglycemia. Fifth, in a previous study, the intake of salt and water affected nocturia^{31,32}, but we could not estimate the intake of salt and water in the present study. Nocturia is likely a multifactorial disorder, and we could not rule out residual confounding factors. Finally, we could not control for the patients' socioeconomic status.

In conclusion, we found an inverse association between vegetable intake habits and nocturia in Japanese patients with type 2 diabetes mellitus. Further studies are required to show the beneficial intake of vegetables for nocturia among patients with type 2 diabetes mellitus.

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DISCLOSURE

The authors declare no conflict of interest.

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