Prevalence of Nocturia Among Community-dwelling Adults: A Population-based Study in Malaysia

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Abstract

Background: Nocturia is widely prevalent condition with detrimental effects on quality of life and general health. In Malaysia, there is a lack of up-to-date prevalence study on nocturia. This study aimed to investigate the prevalence of nocturia and awareness pertaining to nocturia among Malaysian adults.

Methods: A cross-sectional population-based study was conducted among Malaysian adults aged ≥ 18 years old. The data was collected by mixed mode self-administered questionnaire from May 2019 to September 2019. Nocturia was defined as one or more voids at night.

Results: There were a total of 4616 respondents with 74.5% of response rate. The overall prevalence of nocturia among Malaysian adults was found to be 57.3%. In multivariate analysis, respondents aged 31 – 40 (1.91 [1.52 - 2.40]) or > 60 years old (2.03 [1.48 - 2.71]), and those who presented with hypertension (2.84 [2.28 - 3.53]), diabetes mellitus (1.78 [1.42 - 2.25]), renal disease (3.58 [1.93 - 6.63]) or overactive bladder (1.61 [1.10 - 2.35]) were associated with higher prevalence of nocturia. A significantly lower disease prevalence (p < 0.05) was noted among those aged 41 – 50 (0.73 [0.59 - 0.91]), male (0.78 [0.69 - 0.88]) and Chinese (0.47 [0.30 - 0.74]) or Indian (0.34 [0.21 - 0.54]) ethnicities. A total of 37.3% of respondents with nocturia reported that they faced sleeping difficulty about half the time or more after waking up in the middle of night. Those who had ≥ 2 voids per night experienced significantly higher mean bother score than those who had 1 void per night (p < 0.001). Approximately half (56.7%) of all respondents were not aware that night time urination is a medical condition. Only 25.2% of respondents with nocturia had sought medical attention for their nocturia.

Conclusion: The prevalence of nocturia among Malaysian adults is high and strongly influenced by age, gender, race and comorbidities. However, the general awareness on nocturia among Malaysians is poor and relates to low treatment-seeking behaviour we found. The findings also highlighted the impact of nocturia on sleep and the need for nocturia education to better address this disease.

Background

Nocturia is defined as ‘the complaint that the individual has to wake at night one or more times to void’ by International Continence Society (ICS) [1]. Until recently, the definition has been revised as ‘waking up to pass urine during the main sleep period’ [2]. Nocturia can affect all age groups although the disease prevalence does increase with age. Bosch and Weiss reviewed the nocturia prevalence studies in community based, predominantly populations in the western countries and some Asian countries (including China, Taiwan, Japan, Korea, Hong Kong and Singapore)[3]. They reported that prevalence of nocturia among younger populations (20 – 40 years old) with one or more voids per night varies from 11 – 44% whilst those with two or more voids per night has been reported to range between 2 – 18% (Bosch and Weiss 2010) [3]. Nevertheless, nocturia is more frequent among the older populations (> 70 years old), with 69 – 93% having one or more voids per night while 28 – 62% reported two or more voids per night [3]. Similarly, a longitudinal study conducted by van Doorn and colleagues in the Dutch municipality
Krimpen aan den IJssel showed that the prevalence of nocturia increases over time and with increasing age among men aged 50 to 78 years [4].

Several studies have demonstrated that nocturia has negative impact on quality of life and general health [5-8]. Increase in the number of voids per night has serious consequences on sleep, symptoms, morbidity from fatigue and falls and health-related quality of life [5-7]. Unfortunately, nocturia is often unreported and most of those who are affected do not seek medical consultation due to the lack of knowledge that nocturia is treatable [9].

In Malaysia, there is lack of up-to-date prevalence studies on nocturia, only very limited studies are available as reference. A cross-sectional study had been carried out in Institute of Urology and Nephrology (IUN), Kuala Lumpur, Malaysia in 2007 and reported that the prevalence of nocturia was 29% among patients presented with lower urinary tract symptoms (LUTS) [10]. Similarly, Low and colleagues reported that high prevalence (19%) of LUTS was found among women in Northern Malaysia in 2006 and many of them did not seek treatment due to ignorance [11]. Most of these studies were at least 10 years ago, and thus the relevance remains questionable. Therefore, the aim of this present population-based study was to investigate the prevalence of nocturia and awareness pertaining to nocturia among community-dwelling adults in Malaysia.

Methods

A cross-sectional population-based study was conducted among community-dwelling Malaysian adults aged ≥ 18 years old across all states in Malaysia from May 2019 to September 2019 (Figure 1). Convenience sampling was applied to obtain responses from any Malaysian adult aged ≥ 18 year old. The respondents were approached based on the age population percentage reported by Department of Statistics Malaysia (https://www.dosm.gov.my). Individuals who were unable to read or understand English, Malay or Chinese, or with cognitive impairment or active mental illness, and all incomplete responses were excluded. A complete response was considered when the respondent answered all required questions until the last page of the survey, whereas for the questionnaire that was not done until the last page of survey was categorised as ‘incomplete response’.

Questionnaire

The questionnaire consisted of three sections was developed for this study. The first section of the research questionnaire consisted of items to extract the demographic data of the subjects. The second part was made up of three domains, namely present medical condition and disease pattern (2 items), night urination and sleeping issues (5 items) and treatment-seeking behaviour (2 items). Nocturia was defined as one or more voids at night based on the definition from ICS [1]. Those who had to wake up from sleep at night to urinate at least once were required to answer all items, while those without night urination were exempted from some questions pertaining to night urination and sleeping issues and treatment-seeking behaviour. The questionnaire was validated by a panel of experts on content validity and clarity. The English version of the questionnaire was checked by a native English speaker. The
questionnaire was translated into Malay and Chinese languages based on ISPOR guideline with two forward-translation and two back-translation [12]. Three languages of questionnaire were pre-tested with a pilot study (n = 40, respectively) to confirm on the simplicity of language used and to assess the comprehension of the questions. Feedbacks were taken into consideration to incorporate into the final questionnaire. A second pilot study was conducted after the changes made to the questions, followed by a retest of the pilot study to assess internal consistency.

Data Collection

Previous to the data collection, the researchers or the trained research assistants used a pre-trained structured protocol to introduce the instrument to the participants based on the inclusion criteria. In order to increase the response rate, data was collected by mixed mode, i.e. face-to-face and online modes. Face-to-face data collection took place in public areas i.e. shopping malls, parks, restaurants, cafeteria and markets in rural and urban areas. The approached subjects spent approximately 5-10 minutes on the self-administered questionnaires. For the online mode, a third party online platform was utilised to collect the responses. An information sheet about this research study along with a written consent form were attached to each copy of the questionnaire. All subjects signed the written informed consent form prior to answering the questionnaire.

Data Analysis

Continuous variables were presented as mean and standard deviation; whilst categorical variables were presented as frequency and percentage. All data collected were tabulated and analysed using the Statistical Package for Social Sciences software, version 24.0. Pearson's chi-squared ($\chi^2$) test was used to assess the significance in prevalence differences between dichotomous variables, such as gender, age group and employment status (or Fisher's exact test when the frequency of respondents was less than 5 for any category). For ordinal or continuous variables (e.g. age and degree of bother), the Mann-Whitney U test was used to determine the difference between two independent groups (or Kruskal Wallis test if there was more than two independent groups). Univariable analysis were performed for the respondents' characteristics in relation to prevalence of nocturia using logistic regression analysis. Variables with an association of p <0.05 were selected for multivariable analysis using logistic regression analysis. Significance was defined as p <0.05.

Results

The response rate was 74.5%, meaning that a total of 6198 respondents across the states in Malaysia participated in this study. There were 1538 of incomplete responses and 44 duplication of responses, which were excluded from the study. The demographic data of the study population was shown in Table 1. The comparison of demographic characteristics of study population with and without nocturia was shown in Table 2. Statistical analysis revealed that there was significant difference in prevalence between mean age, genders, race employment status and comorbidities (p < 0.05, respectively).
The overall prevalence of nocturia (at least one void per night) among Malaysian adults was 57.3% (Table 2), with 34.1%, 15.3% and 7.9% having an average of one void per night, two voids per night and three or more voids per night respectively. Figure 2 illustrated the increasing trend of prevalence with age in both gender and frequency of nocturnal voids was associated significantly with gender in different age groups. Younger (18 – 40 years old) and older women (> 60 years old) were more likely to have one void per night than men from the same age group (p < 0.05). Whereas, middle-aged men (41 – 60 years old) were more likely to have two voids per night as compared to the middle-aged women (p < 0.05; Figure 2).

The variables univariately associated with nocturia are presented in Table 3. Age group, gender, race, employment status and comorbidities such as hypertension, diabetes mellitus, renal disease and overactive bladder were significantly associated with the prevalence of nocturia. In the multivariate analysis, age group between 31 – 40 or > 60 years old, and those who were presented with comorbidities included hypertension, diabetes mellitus, renal disease or overactive bladder were associated with significantly higher prevalence of nocturia, whereas age group of 41 – 50, male and Chinese or Indian were associated with a decreased prevalence (Table 3). This study also revealed that the prevalence of nocturia was significantly increased with the drink consumption in the late evening, including large volume of water, tea and coffee (Figure 3; p < 0.001) although evening drinking habit was only one of the few factors predisposing to nocturia since the prevalence of nocturia was also significantly high among those without any drink consumption in the late evening (p < 0.001).

From the sub analyses, the frequency of nocturnal voids among respondents with nocturia and among different age groups has been depicted in Figure 4. Those who were more than 60 years old were more likely to experience more than one nocturnal void as compared to their younger counterparts (Figure 4). The mean age for those with once, twice and three or more times of nocturnal voids was 35.51 ± 14.03, 43.41 ± 14.49 and 46.55 ± 14.97, respectively. The association was significant, which indicated that frequency of nocturnal void increased in tandem with age (p < 0.001).

The disease pattern of nocturia encountered by the respondents with nocturia has been summarised in Table 4. Post urination dribble was the only symptom that differed significantly in male respondents with nocturia compared to female (p < 0.001). Those with higher age (> 50 years old) were more likely to present with almost all the symptoms (p < 0.05).

In terms of impact of nocturia on sleep disturbance, 37.3% of respondents with nocturia reported that they faced sleeping difficulty about half the time or more after waking up in the middle of night, particularly among those with ≥ 2 voids per night (66.5%), aged > 50 (40.6%) and employed for wages (62.3%) (Table 5; p < 0.001). As consequences of sleep disturbance due to nocturia, they complaint of fatigue (60.9%), difficulty in concentrating (43.2%), bad mood (41.2%), lack of motivation to work (34.5%), forgetfulness (24.6%), depression (15.6%), lack of sex drive (4.61%) and erectile dysfunction (3.5%; only male respondents). On a scale of 1 to 10 for degree of bother, those who had two voids as well as ≥ 3
voids per night experienced significantly higher mean bother score (3.96 ± 2.41 and 3.91 ± 2.20, respectively) than those who had one void per night (3.29 ± 2.63) (p < 0.001).

Our study also found that the awareness on nocturia was poor among study population. Out of 4616 respondents, 56.7% of them were not aware that night time urination is a medical condition. Unfortunately, the majority (62.2%) of respondents with nocturia did not think that it is a medical condition. Only a minority (25.2%) of respondents with nocturia had sought medical attention for their nocturia (Figure 5). There was a significant association between treatment-seeking behaviour and awareness on nocturia among respondents with nocturia (p < 0.001), which indicated that those who had not sought treatment were not aware that nocturia is a medical condition (64.6%) as compared to those who were aware of this (35.4%). A total of 60.7% out of 4616 respondents had sought or would consider seeking medical attention for nocturia from urologist. There were 39.6% and 21.3% who opted for general practitioner and gynecologist, respectively. This was followed by 17.7% and 17.2% who had chosen pharmacist and nephrologist, respectively.

Discussion

This study presented the first nation-wide report on prevalence of nocturia (at least one void at night) in a multiracial population of Malaysia. The overall prevalence of nocturia among Malaysian adults was high (57%), which was comparable with those reported from studies in Asian population, yet relatively higher compared to Western populations. Liew and colleagues reported that the prevalence of nocturia among Singaporean adults was 55% [13]. Meanwhile, the reported prevalence were 46% in the Dutch population [14], 31% in the US population [5] and 28.4% in the Turkish population [15]. In the present study, there was a significant higher prevalence among women. This finding is consistent with the report from previous studies [13,14,16,17]. Nevertheless, information on the exact etiology pertaining to high prevalence of nocturia in women is lacking. Nocturia was found to be more common among younger women (18 – 40 years old). This is congruent with the finding from study in Finland [17].

In addition, the prevalence of nocturia was found to increase with age, which corroborated the findings of other studies [4,5,18]. This may be due to various pathophysiological mechanisms that related to ageing. In the elderly, nocturia can be resulted from decline in functional bladder capacity with high post void residual, deregulation of arginine vasopressin level, sleep disorders and comorbidities including diabetes mellitus and diabetes insipidus [19]. This also further explains the finding in current study that the frequency of nocturnal voids was higher among the elderly population than younger population.

Previous studies have reported the differences in prevalence of nocturia among races in the US population. The prevalence of nocturia was reported to be higher among Black and Hispanic respondents compared to the White respondents [20,21]. Consistently, similar observations was reported among the
multi-ethnic groups in Malaysia. Malay respondents had a significant higher prevalence compared to other races as revealed by the univariate analysis, yet this prevalence was not significant in multivariate analysis. Chinese and Indian respondents on the other hand, had significantly lower prevalence compared to Malay in both univariate and multivariate analyses. However, Liew and colleagues reported that there was no significant difference between Malay, Chinese and Indian respondents in Singapore [13]. Kupelian and colleagues reported that socioeconomic status was accounted for the ethnic discrepancies in prevalence of nocturia [21]. Further investigation is required in order to identify the underlying etiology of ethnic discrepancies among Malaysians in term of prevalence of nocturia.

Nocturia is strongly associated with various risk factors and comorbidities, including cardiovascular diseases, diabetes, sleep disorders, somatic and mental health [6,13,18,22,23]. This supports the findings of this study which also showed the significant higher disease prevalence among respondents with comorbidities such as hypertension, diabetes mellitus, renal disease or overactive bladder. The pathophysiological changes in hypertensive patients appear to be a significant risk factor for nocturia [24,25]. A prolonged hyperglycemic state in diabetes mellitus patients on the other hand, may cause an alteration in urinary bladder activity, predisposing to diabetic cystopathy [26]. Diabetic cystopathy is primarily induced by paralysis of detrusor muscle and hyperglycemia-induced polyuria that can further lead to bladder hypertrophy in patients with diabetes mellitus [26]. Moreover, microvascular complications of diabetes, particularly diabetic nephropathy and neuropathy have been implicated in nocturia [27]. This explains the mechanistic link between diabetes mellitus and nocturia.

Several studies have reported that nocturia is associated with renal insufficiency [13,28,29]. This may be attributed to the failure of kidneys to concentrate the urine as the renal function worsens, and ultimately leads to nocturnal polyuria and increased nocturnal urine volume in patients with chronic kidney disease [30]. Additionally, overactive bladder was also found to be one of the risk factors in this study. Indeed, nocturia is a common urinary symptoms in overactive bladder patients [31]. However, in this study, there was no significant association between nocturia and benign prostatic hyperplasia, prostate cancer, bladder cancer and snoring, which had been reported in previous studies [22,23,32]. This could be due to the comorbidities determined by self-reporting of respondents in the present study, which might have led to underestimation of the prevalence of these comorbidities in Malaysia.

Excessive evening intake of fluid, caffeinated drinks or alcohol is known to be one of the causes of nocturnal polyuria [1]. In the current study, approximately one third of the respondents with nocturia have the habit of consuming large volume of water, tea or coffee in the late evening. This may suggest that limiting fluid or caffeinated drink intake in the late evening among Malaysian could be one of the modifiable lifestyle risk factors for nocturia. Shiri and colleagues reported that there was no significant association between coffee consumption and nocturia among 50 to 70 years old men in Finland, yet the risk of moderate or severe nocturia was 2-fold higher among men who drank 6 cups coffee or more per day than non-drinkers or those who consumed one cup per day for this age group [33]. Interestingly, the
prevalence of nocturia was about one third of respondent despite any drink consumption. The actual prevalence of nocturia with underlying pathological causes in the population may be reflected by eliminating the potential confounding factor of drink consumption in the late evening.

LUTS can be classified according to voiding or obstructive symptoms (such as hesitancy, poor stream, straining, prolonged micturition, feeling of incomplete bladder emptying, post-micturition dribble) and storage or irritative symptoms (such as frequency, urgency, urge incontinence and nocturia). In the present study, the disease pattern of nocturia was not gender-specific, except for post-micturition dribble. This interpretation is supported by the fact that post-micturition dribble is more common in men [34]. Studies also showed that prevalence of LUTS was comparable between men and women [35,36]. In addition, with advancing age, the prevalence and severity of LUTS increase regardless of gender [35]. This is supported by the finding of this study where elderly respondents are more likely to exhibit almost all the LUTS compared to the younger population.

Nocturia has been regarded as not only ‘bothersome’, but also negatively affects sleep and quality of life [5-8]. This study also revealed that nocturia was regarded as moderately bothersome for Malaysians, particularly those with two or more nocturnal voids. Furthermore, more than one-third of respondents in this study claimed to have disturbed sleep about half the time or more after nocturnal awakening. The sleep disturbance was proportionately increased with the frequency of nocturnal voids and age. This finding is consistent with previous studies [5,13,14,37]. As a result of sleep disturbance, nocturia has also been associated with excessive daytime sleepiness (7 – 18%), increased sick leaves (27 – 27%), insomnia (43%), psychiatric disorders (28 – 35%) and poorer health status (15 – 40%) [38,39] In addition, people with nocturia have a lower productivity at work and decreased activity levels [6]. Consistently, the respondents with nocturia in this study also reported to experience fatigue, difficulty in concentrating, lacking of motivation to work, bad mood, forgetfulness and depression as consequences of disturbed sleep. Hence, it is obvious that nocturia should not be overlooked or underestimated due to its serious repercussions on sleep quality, work productivity and quality of life.

Treatment-seeking behaviour for LUTS is correlated with the symptom severity, degree of bother, costs and benefits of treatment [40,41]. To the best of our knowledge, this is the first study to report on the association between awareness of nocturia and treatment-seeking behaviour in Malaysia. Approximately two-thirds of respondents with nocturia in this study were not aware that nocturia is a medical condition and this was significantly associated with behaviour of not seeking medical attention. This finding suggests that poor awareness might be a factor affecting their treatment-seeking behaviour. In Taiwan, Chen and colleagues found that the lack of knowledge on nocturia appeared to be an important barrier to medical consultation among Taiwanese women [9]. In addition, a vast majority of those who had not sought treatment in the present study perceived that nocturia was not a significant medical issue to seek treatment. This appear consistent with finding by Low and colleagues who reported that majority of women in Northern Malaysia did not seek treatment for LUTS due to ignorance [11]. These findings could
be attributed to poor awareness and knowledge pertaining to nocturia among Malaysians. As such, addressing the misconceptions about nocturia can help to increase the awareness, enhance treatment-seeking behaviour and also improve the overall quality of life among people with nocturia.

In the present study, convenience sampling was conducted instead of stratified sampling to define the population in Malaysia. Despite this limitation, the subjects sampled were closely reflecting the real population in Malaysia. Although there was slightly difference between the sample and population in terms of age and gender, these differences were adjusted in the statistical analysis. Hence, the outcomes of this study are valuable to provide sights into prevalence of nocturia among Malaysians.

**Conclusion**

The prevalence of nocturia among Malaysian adults is high and strongly influenced by age, gender, race and comorbidities. There is a significant correlation between frequency of nocturnal voids with sleep disturbance and degree of bother. However, the general awareness on nocturia was poor among Malaysians. Our study provides insights into the factors affecting the treatment-seeking behaviour for nocturia. The behaviour of not-seeking medical attention was associated to poor awareness and misconception on nocturia in the Malaysian population. The findings serve as an important feedback to all stakeholders to better address this disease.

**Abbreviations**

ICS: International Continence Society; LUTS: lower urinary tract symptoms; S.D.: standard deviation; OR: Odd ratio; CI: Confidence interval

**Declarations**

**Ethics approval and consent to participate**

Ethical approval was obtained from the Human Ethics Committee of Taylor’s University, Malaysia (HEC 2019/012). An information sheet about this research study along with a written consent form were attached to each copy of the questionnaire. All subjects signed the written informed consent form prior to answering the questionnaire.

**Consent for publication**

Subject consent was taken for publication on data and identity confidentiality. As the data were reported in a collective manner with no reference to an individual, the subject’s identity could be kept confidential.

**Availability of data and materials**

The datasets used and/or analysed during the current study are available from the corresponding author on reasonable request. The English version of questionnaire is available in the supplementary
information file.

**Competing interests**

All authors declare that they have no competing interests.

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**Authors' contributions**

We declare that this work was done by the authors named in this article and all liabilities pertaining to claims relating to the content of this article will be borne by them. YHY, JTJL, MCW, OTA, ZMZ, TCC, NKS, GEH and ANF developed the idea to conduct this research. YHY, EWF, JTJL and MCW designed and planned the research and pilot study, as well as performed the literature search. YHY, JTJL, MCW and LSH conducted the data collection, YHY and MCW performed the data and statistical analyses. The manuscript was written and edited by YHY, and reviewed by all authors.

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**References**

1. van Kerrebroeck P, Abrams P, Chaikin D, Donovan J, Fonda D, Jackson S, et al. The standardisation of terminology in nocturia: Report from the Standardisation Sub-committee of the International Continence Society. BJU Int. 2002;90:11-5.

2. International Continence Society. Terminology discussion: Nocturia. 2018. [https://www.ics.org/committees/standardisation/terminologydiscussions/nocturia](https://www.ics.org/committees/standardisation/terminologydiscussions/nocturia). Accessed 17...
3. Bosch JL, Weiss JP. The Prevalence and Causes of Nocturia. J Urol. 2010;184:440-6.
4. van Doorn B, Blanker MH, Kok, ET, Westers P, Bosch JL. Once nocturia, always nocturia? Natural history of nocturia in older men based on frequency-volume charts: the Krimpen study. J Urol. 2011;186:1956-61.
5. Coyne KS, Zhou Z, Bhattacharyya SK, Thompson CL, Dhawan R, Versi E. The prevalence of nocturia and its effect on health-related quality of life and sleep in a community sample in the USA. BJU Int. 2003;92:948-54.
6. Asplund R, Marnetoft SU, Selander J, Akerström B. Nocturia in relation to somatic health, mental health and pain in adult men and women. BJU Int. 2005;95:816-9.
7. Weiss JP, Blaivas JG, Blanker MH, Bliwise DL, Dmochowski RR, Drake M, et al. The New England Research Institutes, Inc. (NERI) Nocturia Advisory Conference 2012: focus on outcomes of therapy. BJU Int. 2013;700-16.
8. Kupelian V, Fitzgerald MP, Kaplan SA, Norgaard JP, Chiu GR, Rosen RC. Association of nocturia and mortality: results from the Third National Health and Nutrition Examination Survey. J Urol. 2011;185:571-7.
9. Chen FY, Dai YT, Liu CK, Yu HJ, Liu CY, Chen TH. Perception of nocturia and medical consulting behavior among community-dwelling women. Int Urogynecol J Pelvic Floor Dysfunct. 2007;18:431-6.
10. Mariappan P, Turner KJ, Sothilingam S, Rajan P, Sundram M, Stewart LH. Nocturia, nocturia indices and variables from frequency-volume charts are significantly different in Asian and Caucasian men with lower urinary tract symptoms: a prospective comparison study. BJU Int. 2007;100:332-6.
11. Low BY, Liong ML, Yuen KH, Chong WL, Chee C, Leong WS, et al. Study of prevalence, treatment-seeking behavior, and risk factors of women with lower urinary tract symptoms in Northern Malaysia. Urology. 2006;68:751-8.
12. Wild D, Eremenco S, Mear I, Martin M, Houchin C, Gawlicki M, et al. Multinational trials – recommendations on the translations required, approaches to using the same language in different countries, and the approaches to support pooling the data: the ISPOR Patient Reported Outcomes Translation & Linguistic Validation Good Research Practices Task Force Report. Value Health. 2009;12:430-40.
13. Liew LC, Tiong HY, Wong ML, Png DC, Tan JK. A population study of nocturia in Singapore. BJU Int. 2006;97:109-12.
14. van Dijk L, Kooij DG, Schellevis FG. Nocturia in the Dutch adult population. BJU Int. 2002;90:644-8.
15. Zumrutbas AE, Bozkurt Al, Alkis O, Toktas C, Cetinel B, Aybek Z. The prevalence of nocturia and nocturnal polyuria: Can new cutoff values be suggested according to age and sex?. Int Neurourol J. 2016;20:304-10.
16. Choo MS, Ku JH, Park CH, Lee YS, Lee KS, Lee JG, et al. Prevalence of nocturia in a Korean population aged 40 to 89 years. Neurourol Urodyn. 2008;27:60-4.
17. Tikkinen KA, Tammela TL, Huhtala H, Auvinen A. Is nocturia equally common among men and women? A population based study in Finland. J Urol. 2006;175:596-600.

18. Kim SY, Bang W, Kim MS, Park B, Kim JH, Choi HG. Analysis of the prevalence and factors associated with nocturia in adult Korean men. Sci Rep. 2017;7:41714.

19. van Doorn B, Bosch JL. Nocturia in older men. Maturitas. 2012;71:8-12.

20. Fitzgerald MP, Litman HJ, Link CL, McKinlay JB. The association of nocturia with cardiac disease, diabetes, body mass index, age and diuretic use: results from the BACH survey. J Urol. 2007;177:1385-9.

21. Kupelian V, Link CL, Hall SA, McKinlay JB. Are racial/ethnic disparities in the prevalence of nocturia due to socioeconomic status? Results of the BACH survey. J Urol. 2009;181:1756-63.

22. Madhu C, Coyne K, Hashim H, Chapple C, Milsom I, Kopp Z. Nocturia: risk factors and associated comorbidities; findings from the EpiLUTS study. Int. J. Clin. Pract. 2015;69:1508-16.

23. Yoshimura K, Terada N, Matsui Y, Terai A, Kinukawa N, Arai Y. Prevalence of and risk factors for nocturia: analysis of a health screening program. Int J Urol. 2004;11:282-7.

24. Feldstein CA. Nocturia in arterial hypertension: a prevalent, underreported, and sometimes underestimated association. J. Am. Soc. Hypertens. 2013;7:75-84.

25. McKeigue PM, Reynard JM. Relation of nocturnal polyuria of the elderly to essential hypertension. Lancet 2000;355:486-8.

26. Barua SK, Borah SJ, Rajeev TP. Current understanding of diabetes mellitus-induced lower urinary tract dysfunction. J Health Res Rev. 2019;6:50-6.

27. Hill SR, Fayyad AM, Jones GR. Diabetes mellitus and female lower urinary tract symptoms: a review. Neurourol. Urodyn. 2008;27:362-7.

28. Fujimura T, Yamada Y, Sugihara T, Azuma T, Suzuki M, Nakagawa T, et al. Nocturia in men is a chaotic condition dominated by nocturnal polyuria. Int J Urol. 2015;22:496-501.

29. Wu MY, Wu YL, Hsu YH, Lin YF, Fan YC, Lin YC, et al. Risk of nocturia in patients with chronic kidney disease—do the metabolic syndrome and its components matter?. J Urol. 2012;188:269-73.

30. Fukuda M, Motokawa M, Miyagi S, Sengo K, Muramatsu W, Kato N, et al. Polynocturia in chronic kidney disease is related to natriuresis rather than to water diuresis. Nephrol Dial Transplant. 2006;21:2172-7.

31. Espuña-Pons M, Blasco P, Pérez M, Rebollo P. Nocturia in patients with overactive bladder. Arch Esp Urol. 2010;63:363-72.

32. Tikkinen KA, Auvinen A, Johnson TM 2nd, Weiss JP, Keranen T, Tiitinen A, et al. A systematic evaluation of factors associated with nocturia—The population-based FINNO study. Am J Epidemiol. 2009;170:361-8.

33. Shiri R, Hakama M, Häkkinen J, Auvinen A, Huhtala H, Tammela TLJ, et al. The effects of lifestyle factors on the incidence of nocturia. J Urol. 2008;180:2059-62.
34. Yang DY, Lee WK. A current perspective on post-micturition dribble in males. Investig Clin Urol. 2019;60:142-7.

35. Boyle P, Robertson C, Mazzetta C, Keech M, Hobbs FDR, Fourcade R, et al. The prevalence of lower urinary tract symptoms in men and women in four centres. The UrEpiK study. BJU Int. 2003;92:409-14.

36. Diokno AC, Brock BM, Brown MB, Herzog AR. Prevalence of urinary incontinence and other urological symptoms in the noninstitutionalized elderly. J Urol. 1986;136:1022-5.

37. Kim SO, Choi HS, Kim YJ, Kim HS, Hwang IS, Hwang EC, et al. Impact of nocturia on health-related quality of life and medical outcomes study sleep score in men. Int Neurourol J. 2011;15:82-6.

38. Ohayon MM. Nocturnal awakenings and comorbid disorders in the American general population. J. Psychiatr. Res. 2008;43:48-54.

39. Asplund R, Aberg H. Health of the elderly with regard to sleep and nocturnal micturition. Scan J Prim Health Care. 1992;10:98-104.

40. Welliver C, Sulaver R, Whittington A, Helfand BT, Çakır OO, Griffith JW, et al. Analyzing why men seek treatment for lower urinary tract symptoms and factors associated with nonimprovement. Urology. 2015;86:862-7.

41. Griffith JW, Messersmith EE, Gillespie BW, Wiseman JB, Flynn KE, Kirkali Z, et al. Reasons for seeking clinical care for lower urinary tract symptoms: A mixed methods study. J Urol. 2018;199:528-35.

Tables

Table 1: Demographic characteristic of study population
| Variable                        | Overall (n,%) | Mode of questionnaire | P value* |
|--------------------------------|---------------|-----------------------|---------|
|                                | Online (n, %) | Face-to-Face (n, %)   |         |
| Total respondents              | 4616 (100)    | 4004 (86.7)           | 612 (13.3) |
| Mean age (years ± S.D.)        | 38.4 ± 14.58  | 38.0 ± 14.08          | 40.9 ± 17.31 | 0.025 φ |
| 18 – 30                        | 1794 (38.9)   | 1573 (39.2)           | 221 (36.2) |
| 31 – 40                        | 889 (19.3)    | 835 (20.9)            | 54 (8.8)  |
| 41 – 50                        | 760 (16.5)    | 692 (17.3)            | 68 (11.1) |
| 51 – 60                        | 865 (18.7)    | 664 (16.6)            | 201 (32.9)|
| > 60                           | 307 (6.7)     | 240 (6.0)             | 67 (11.0) |
| Gender                         |               |                       |         |
| Male                           | 1982 (42.9)   | 1646 (41.1)           | 336 (54.9) | <0.001 # |
| Female                         | 2634 (57.1)   | 2358 (58.9)           | 276 (45.1)|          |
| Race                           |               |                       |         |
| Malay                          | 1796 (38.9)   | 1222 (30.5)           | 574 (94.1) | <0.001 # |
| Chinese                        | 2126 (46.1)   | 2115 (52.8)           | 11 (1.8)  |
| Indian                         | 595 (12.9)    | 574 (14.4)            | 21 (3.4)  |
| Others                         | 96 (2.1)      | 92 (2.3)              | 4 (0.7)   |
| Employment Status              |               |                       |         |
| Student                        | 920 (19.9)    | 724 (18.1)            | 196 (32.0) | <0.001 # |
| Employed for wages             | 2659 (57.4)   | 2370 (59.2)           | 280 (45.8)|          |
| Self-employed                  | 394 (8.5)     | 319 (8.0)             | 75 (12.2) |
| Unemployed                     | 652 (14.1)    | 591 (14.7)            | 61 (10.0) |
| Comorbidities                  |               |                       |         |
| Hypertension                   | 638 (13.8)    | 502 (12.5)            | 136 (22.2) | <0.001 # |
| Diabetes mellitus              | 524 (11.4)    | 402 (10.0)            | 122 (19.9) | <0.001 # |
| Renal disease                  | 90 (1.9)      | 39 (1.0)              | 51 (8.3)  | <0.001 # |
| Heart failure                  | 22 (0.5)      | 16 (0.4)              | 6 (1.0)   |
| Enlarged prostate              | 61 (1.3)      | 59 (1.5)              | 2 (0.3)   | 0.052 # |
| Overactive bladder             | 141 (3.1)     | 121 (3.0)             | 20 (3.3)  | 0.021 # |
|                      | n (%)          | n (%)          | n (%)          | p-value |
|----------------------|----------------|----------------|----------------|---------|
| Prostate cancer      | 9 (0.2)        | 8 (0.2)        | 1 (0.2)        | 0.742*  |
| Bladder cancer       | 6 (0.1)        | 6 (0.1)        | 0 (0)          | 1.000φ  |
| Snoring              | 781 (16.9)     | 714 (91.4)     | 67 (11)        | <0.001# |
| Prevalence of Nocturia | 2646 (57.3) | 2166 (54.1)    | 480 (78.4)     | <0.001# |

*The statistical significance level was 0.05; φMann-Whitney U test; #Chi-square test; ϕFisher’s Exact test; S.D.: standard deviation; n = 4616.

**Table 2:** Demographic characteristic of study population with and without nocturia
| Variable                        | Nocturia (n, %) | No Nocturia (n, %) | P value* |
|--------------------------------|-----------------|--------------------|----------|
| Total respondents              | 2646 (57.3)     | 1970 (42.7)        |          |
| Mean age (years ± SD)          | 39.14 ± 14.98   | 37.33 ± 13.96      | < 0.001  |
| 18 – 30                        | 948 (35.8)      | 846 (43.0)         |          |
| 31 – 40                        | 591 (22.4)      | 298 (15.1)         |          |
| 41 – 50                        | 366 (13.8)      | 394 (20.0)         |          |
| 51 – 60                        | 508 (19.2)      | 357 (18.1)         |          |
| > 60                           | 233 (8.8)       | 74 (3.8)           |          |
| Gender                         |                 |                    |          |
| Male                           | 1103 (41.7)     | 879 (44.6)         | 0.046#   |
| Female                         | 1543 (58.3)     | 1091 (55.4)        |          |
| Race                           |                 |                    |          |
| Malay                          | 1239 (46.9)     | 557 (28.2)         | < 0.001# |
| Chinese                        | 1087 (41.1)     | 1039 (52.8)        |          |
| Indian                         | 253 (9.6)       | 342 (17.4)         |          |
| Others                         | 65 (2.4)        | 31 (1.6)           |          |
| Employment Status              |                 |                    |          |
| Student                        | 490 (18.5)      | 430 (21.8)         | 0.019#   |
| Employed for wages             | 1555 (58.8)     | 1095 (55.6)        |          |
| Self-employed                  | 216 (8.2)       | 178 (9.0)          |          |
| Unemployed                     | 385 (14.5)      | 267 (13.6)         |          |
| Comorbidities                  |                 |                    |          |
| Hypertension                   | 500 (18.90)     | 138 (7.0)          | < 0.001# |
| Diabetes mellitus              | 369 (13.95)     | 155 (7.9)          | < 0.001# |
| Renal disease                  | 77 (2.91)       | 13 (0.7)           | < 0.001# |
| Heart failure                  | 17 (0.64)       | 5 (0.3)            |          |
| Enlarged prostate              | 41 (1.55)       | 20 (1.0)           | 0.058#   |
| Overactive bladder             | 93 (3.51)       | 48 (2.4)           | 0.116#   |
| Prostate cancer                | 6 (0.23)        | 3 (0.2)            | 0.035#   |
|                        | No Nocturia | Nocturia | p-value |
|------------------------|-------------|----------|---------|
| Bladder cancer         | 4 (0.15)    | 2 (0.1)  | 0.741φ  |
| Snoring                | 471 (17.80) | 310 (15.7)| 1.000φ  |

*The statistical significance level was 0.05; φMann-Whitney U test; #Chi-square test; ϕFisher’s Exact test; S.D.: standard deviation; n = 4616.

**Table 3:** Univariable and multivariable analysis for respondents’ characteristic in relation to prevalence of nocturia
|                | Univariate Analysis* |                       | Multivariate Analysis*# |                       |
|----------------|----------------------|-----------------------|-------------------------|-----------------------|
|                | OR (95% CI)          | p value               | OR (95% CI)             | p value               |
| Age group      |                      |                       |                         |                       |
| 18 – 30        | 0.741 (0.658 – 0.835)| <0.001                | 1.161 (0.931 – 1.449)   | 0.185                 |
| 31 – 40        | 1.613 (1.383 – 1.880)| <0.001                | 1.909 (1.521 – 2.396)   | <0.001                |
| 41 – 50        | 0.642 (0.549 – 0.750)| <0.001                | 0.733 (0.592 – 0.908)   | 0.005                 |
| 51 – 60        | 1.073 (0.923 – 1.803)| 0.358                 | Not included            | -                     |
| > 60           | 2.473 (1.891 – 3.234)| <0.001                | 2.027 (1.478 – 2.708)   | <0.001                |
| Gender         |                      |                       |                         |                       |
| Male           | 0.887 (0.789 – 0.998)| 0.046                 | 0.779 (0.687 – 0.884)   | <0.001                |
| Female         | 1.127 (1.002 – 1.268)| 0.046                 | -                       | -                     |
| Race           |                      |                       |                         |                       |
| Malay          | 2.235 (1.974 – 2.531)| <0.001                | 0.891 (0.569 – 1.396)   | 0.615                 |
| Chinese        | 0.625 (0.556 – 0.703)| <0.001                | 0.471 (0.302 – 0.735)   | 0.001                 |
| Indian         | 0.503 (0.423 – 0.599)| <0.001                | 0.336 (0.210 – 0.537)   | <0.001                |
| Others         | 1.576 (1.023 – 2.427)| 0.039                 | -                       | -                     |
| Employment status |                     |                       |                         |                       |
| Student        |                      |                       |                         |                       |
| Employed for wages | 0.814 (0.704 – 0.941)| 0.005                | 1.154 (0.911 – 1.463)   | 0.235                 |
| Self-employed  | 1.139 (1.012 – 1.281)| 0.031                 | 1.141 (0.963 – 1.352)   | 0.129                 |
| Unemployed     | 0.895 (0.727 – 1.101)| 0.294                 | Not included            | -                     |
|                | 1.086 (0.918 – 1.285)| 0.336                 | Not included            | -                     |
| Comorbidities  |                      |                       |                         |                       |
| Hypertension   | 3.093 (2.536 – 3.772)| <0.001                | 2.837 (2.281 – 3.530)   | <0.001                |
| Diabetes mellitus | 1.898 (1.558 – 2.312)| <0.001                | 1.783 (1.416 – 2.245)   | <0.001                |
| Renal disease  |                      |                       |                         |                       |
| Heart failure  | 4.512 (2.500 – 8.145)| <0.001                | 3.579 (1.933 – 6.625)   | <0.001                |
| Enlarged prostate | 2.541 (0.936 – 6.900)| 0.067                 | Not included            | -                     |
| Overactive bladder | 1.535 (0.896 – 2.627)| 0.119                 | Not included            | -                     |
| Disease Pattern          | Odds Ratio (CI)          | p-Value | Odds Ratio (CI)          | p-Value |
|-------------------------|-------------------------|---------|-------------------------|---------|
| Prostate cancer         | 1.459 (1.024 – 2.077)   | 0.036   | 1.606 (1.097 – 2.352)   | 0.015   |
| Bladder cancer          |                         |         |                         |         |
| Snoring                 | 1.490 (0.372 – 5.966)   | 0.573   | Not included            | -       |
|                         | 1.490 (0.273 – 8.142)   | 0.645   | Not included            | -       |
|                         | 1.160 (0.991 – 1.357)   | 0.064   | Not included            | -       |

*Based on logistic regression analysis, the statistical significance level was 0.05; Variables with an association of p <0.05 were selected for a multivariable analysis; OR: Odd ratio; CI: Confidence interval; n = 4616.

**Table 4:** Disease pattern of nocturia among study population based on gender and age
| Symptom                                                        | Gender (n, %) | Age group (n, %) |
|---------------------------------------------------------------|--------------|-----------------|
|                                                               | Male         | Female          | 18 – 50 | > 50     |
| Total respondents                                            | 1103 (41.7)  | 1543 (58.3)     | 1905 (72.0) | 741 (28.0) |
| Post urination dribble                                       | 172 (15.6)   | 139 (9.0)       | 237 (12.4) | 74 (10.0)  |
|                                                               | p < 0.001*   | p = 0.078*      |          |          |
| Frequent urination during the day                            | 276 (25.0)   | 274 (17.8)      | 438 (23.0) | 212 (28.6) |
|                                                               | p = 0.644*   | p = 0.003*      |          |          |
| Frequent urinary urgency                                     | 129 (11.7)   | 168 (10.9)      | 155 (8.1) | 142 (19.2) |
|                                                               | p = 0.516*   | p < 0.001*      |          |          |
| Urinary Incontinence                                         | 90 (8.2)     | 124 (8.0)       | 73 (3.8)  | 141 (19.0) |
|                                                               | p = 0.909*   | p < 0.001*      |          |          |
| Slow urine flow                                              | 63 (5.7)     | 78 (5.1)        | 69 (3.6)  | 72 (9.7)  |
|                                                               | p = 0.458*   | p < 0.001*      |          |          |
| Feeling of incomplete bladder emptying after urination       | 143 (13.0)   | 165 (10.7)      | 166 (8.7) | 142 (19.8) |
|                                                               | p = 0.072*   | p < 0.001*      |          |          |

*Based on Chi-square test, the statistical significance level was 0.05; n = 2646.

Table 5: The impact of nocturia on sleep disturbance and factors affecting the sleep disturbance
| Variable                      | Degree of sleep disturbance (n, %) | p value*   |
|-------------------------------|-----------------------------------|------------|
|                               | Less than half the time | About half the time or more |
| Total respondents             | 1655 (62.7)         | 983 (37.3) |          |
| Frequency of Nocturia         |                     |            |          |
| Once                         | 1241 (75.0)         | 329 (33.5) | <0.001   |
| Twice                        | 273 (16.5)          | 433 (44.0) |          |
| Three times or more          | 141 (8.5)           | 221 (22.5) |          |
| Age                          |                     |            |          |
| 18 – 30                      | 730 (44.1)          | 215 (21.9) | <0.001   |
| 31 – 40                      | 375 (22.6)          | 215 (21.9) |          |
| 41 – 50                      | 210 (12.7)          | 154 (15.6) |          |
| 51 – 60                      | 233 (14.1)          | 273 (27.8) |          |
| > 60                         | 107 (6.5)           | 126 (12.8) |          |
| Gender                       |                     |            |          |
| Male                         | 681 (41.1)          | 417 (42.4) | 0.521    |
| Female                       | 974 (58.9)          | 566 (57.6) |          |
| Employment status            |                     |            |          |
| Student                      | 391 (23.6)          | 98 (10.0)  | <0.001   |
| Employed for wages           | 938 (56.7)          | 613 (62.3) |          |
| Self-employed                | 116 (7.0)           | 98 (10.0)  |          |
| Unemployed                   | 210 (12.7)          | 174 (17.7) |          |

*Based on Chi-square test, the statistical significance level was 0.05; n = 2638.

**Figures**
Data collection from May 2019 – September 2019
Inclusion: Malaysian adults aged ≥ 18 years old

Online Questionnaire
n = 5586

Excluded (Total = 1582)
☐ Incomplete responses: n = 1538
☐ Duplication: n = 44

n = 4004

Face-to-face Questionnaire
n = 612

Excluded (Total = 0)

n = 612

Total recruited, n = 4616

Figure 1

Study protocol of subject inclusion and exclusion.
Figure 2

The prevalence of nocturia based on gender and frequency of nocturia. n = 4616. *According to the Chi-Square test, younger (18 – 40 years old) and older women (> 60 years old) were more commonly to have one void per night than men from the same age group (p < 0.05). Whereas, middle-aged men (41 – 60 years old) were more likely to have two voids per night as compared to the middle-aged women (p < 0.05).
Figure 3

Relationship between prevalence of nocturia and drinks consumption in the late evening. $n = 4616$.

*According to the Chi-Square test, the prevalence of nocturia was significantly associated with the drink consumption in the late evening, including large volume of water, tea and coffee ($p < 0.001$). The prevalence of nocturia was significant among those without any drink consumption in the late evening ($p < 0.001$).
Figure 4

Frequency of nocturia (a) among respondents with nocturia, (b) in different age groups. n = 2646.
Figure 5

Treatment seeking behaviour among participants with nocturia. n = 2037. (a) The overall treatment seeking behaviour among participants with nocturia. (b) The responses from those participants who had sought for treatment. (c) The responses from participants who had not sought for treatments.

Supplementary Files
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- QuestionnaireNocturiaEnglishversion.pdf