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## Relationship between Central Obesity, General Obesity, Overactive Bladder Syndrome and Urinary Incontinence Among Male and Female Patients Seeking Care for their Lower Urinary Tract Symptoms

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### Abstract

**Objectives:** To describe the relationship between metabolic factors and lower urinary tract symptoms (LUTS), overactive bladder syndrome (OAB) and urinary incontinence (UI).

**Methods:** Adult male and female patients who presented to a clinician from the Symptoms of Lower Urinary Tract Dysfunction Research Network (LURN) were recruited. Urinary symptoms (presence of OAB, any UI, stress UI (SUI), urgency UI (UUI), urgency, frequency, and nocturia) were assessed with the LUTS Tool. Metabolic factors assessed included central obesity (waist circumference, using the Adult Treatment Panel III, the International Diabetes Federation thresholds, and waist circumference as a continuous variable), general obesity (body mass index [BMI] as dichotomous or continuous variables), diabetes mellitus, hypertension, and dyslipidemia. Multivariable logistic regression was used to test for associations.

**Results:** 920 participants were studied. In multivariable analyses, central obesity (per 10cm larger waist) was associated with higher odds of UI in both sexes (OR=1.16, p=0.008), SUI in females (OR=1.27, p=0.008), UUI in both sexes (OR=1.24, p=0.001), OAB in females (OR=1.248, p=0.003), as well as frequency and nocturia. General obesity (5-unit increase in BMI) was associated with UI, UUI, urgency and frequency in both sexes, and with SUI and OAB in females. We did not find associations between central or general obesity and OAB in males. Dyslipidemia was associated with nocturia 2.

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**Declaration of Interest:** None

**Conclusions:** In patients, central and general obesity were key metabolic factors associated with UI in both males and females, and with OAB in females but not in males. The association between dyslipidemia and nocturia needs further research.

### Keywords

body mass index; central obesity; lower urinary tract symptoms; overactive bladder; urinary incontinence

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### Introduction:

The prevalence of metabolic factors such as obesity and diabetes in the United States has been increasing. The morbidity and mortality associated with metabolic factors are costly and pose a serious burden to society. Recent studies suggest that metabolic factors may play a role in the development of overactive bladder syndrome (OAB), urinary incontinence (UI) and other lower urinary tract symptoms (LUTS). However, the literature has major knowledge gaps. For females, studies have shown high body mass index (BMI) to be a risk factor for UI. However, the relationships between obesity and OAB are less clear. The results are conflicting; some studies showed that high BMI was associated with OAB while other studies showed no association. For males, studies have reported associations between metabolic factors and higher American Urological Association Symptom Index (AUA-SI) scores. However, the relationships between metabolic factors and UI and OAB in males have not been well studied. In addition, most studies were community-based; thus it is unclear whether the findings would apply to patients seeking care for their LUTS. Among studies that have examined obesity, most have only reported on BMI. Few studies have specifically evaluated central obesity for its association with urinary symptoms.

The Symptoms of Lower Urinary Tract Dysfunction Research Network (LURN) sought to address these knowledge gaps by recruiting a large cohort of male and female patients who were seeking care for their LUTS. We assessed multiple metabolic factors (including three measures of central obesity, three measures of BMI, diabetes, hypertension, and dyslipidemia), and their relationships to OAB, UI, and other LUTS. We expect to gain insights into the relevant and modifiable metabolic factor(s) associated with these LUTS that may inform future primary and secondary management and prevention strategies.

### Materials and Methods:

#### Study Design and Population

The LURN Observational Cohort Study recruited adult male and female patients over the age of 18 who presented to a urologist or urogynecologist for treatment of their LUTS between June 2015 and January 2017 at one of six LURN clinical centers in the United States. Participants were followed prospectively for one year, although data reported herein only include baseline data. The inclusion and exclusion criteria and details of the study design have been previously reported. Participants with neurogenic conditions or neurogenic bladder were excluded. The following data were collected at the baseline visit: urinary symptoms, medical history, medication use, systolic and diastolic blood pressure, BMI, and

waist circumference measurements. All participants provided informed consent. The protocol was approved by the institutional review boards of all participating research sites.

## Measures

Central obesity was assessed based on waist circumference measurements. Three measures of central obesity were used: 1) the Adult Treatment Panel III (ATP III) Guidelines of 102 cm for males, 88 cm for females, 2) the 2004 Consensus Statement from the International Diabetes Federation (IDF) of 94 cm for white males, 80 cm for white females. and 3) waist circumference as a continuous variable. Body weight and height were also measured, and BMI was calculated as the weight in kilograms divided by the square of the height in meters. General obesity was defined as BMI  $\geq 30$ ; overweight was defined as BMI  $\geq 25$ ; and BMI was also used as a continuous variable.

The majority of studies in the literature have only assessed BMI. Since central obesity is a key component of metabolic syndrome, we have also included measures of central obesity in addition to BMI. We have included the two most commonly used definitions of central obesity (ATP III and IDF) to be comprehensive. The ATP III definition used the same waist circumference cutoffs regardless of race, while the newer IDF definition used race-specific waist circumference cutoffs. Also, the IDF used lower cutoffs to include subjects who are overweight but not yet considered obese.

Hypertension was defined by a measured systolic pressure  $\geq 130$  mmHg, diastolic pressure  $\geq 85$  mmHg, or self-reported medication(s) to treat hypertension. Dyslipidemia was defined by self-reported medication(s) taken to treat elevated serum cholesterol or triglycerides. Diabetes was defined by a self-reported medical history of diabetes mellitus.

Urinary symptoms were assessed using the LUTS Tool with a one-week recall period. Seven LUTS Tool questions addressed urinary incontinence (UI); participants with responses of “rarely” or “never” to all seven UI questions were classified as “without UI” and responses of “sometimes” or greater to at least one symptom of UI were classified as “with UI”. Two LUTS Tool questions addressed stress urinary incontinence (SUI), leakage while exercising or during a laugh, cough, or sneeze. Participants were classified as having SUI if they had responses of “sometimes” or more to at least one of two leakage questions. Participants who answered “sometimes” or more to a sudden need to rush to urinate were classified as having urgency. A follow-up question asked about urinary leakage due to a sudden feeling of needing to rush to urinate; those who responded “sometimes” or more were classified as having urgency urinary incontinence (UUI).

Participants who reported that they “sometimes”, “often” or “almost always” urinate too frequently were classified as having frequency. Those who reported that they wake up because of the need to urinate one or two times a night or more were classified as having nocturia 1 or nocturia 2 respectively. Participants who answered affirmatively to the urgency question, and who also reported frequency and/or nocturia were classified as having overactive bladder syndrome (OAB), in accordance with the definition of the International Continence Society (ICS). Among those with OAB, they were further classified into two groups: with UUI (also referred as “OAB-wet”) versus without UUI (“OAB-dry”).

## Statistics

Demographics and other participant characteristics are shown as means and standard deviations, or medians and interquartile ranges, as appropriate, for continuous variables and as frequencies and percentages for categorical variables.

Multivariable logistic regression models were fitted to test associations between metabolic factors and the LUTS groups described above. For each outcome (e.g. UI, OAB), regression models were developed using the following metabolic predictors: central obesity cutoffs defined by ATP III or by IDF, waist circumference as a continuous variable, general obesity (BMI  $\geq 30$ ), overweight status (BMI  $\geq 25$ ), BMI as a continuous variable, hypertension, diabetes, and dyslipidemia. Additional candidate covariates included sex, age, race, ethnicity, and smoking status. Menopausal status, hormonal use, parity, and pelvic organ prolapse quantification classification were also included for females using interactions with sex. Model selection was guided by the method of best subsets. Interactions between sex and waist circumference/BMI (both continuous and dichotomous versions) were tested in all models and are presented when statistically significant ( $p < 0.05$ ). All p-values were adjusted for multiple comparisons using the false discovery rate (FDR) correction. Models are shown with the adjusted odds ratios and 95% confidence levels. All analyses were conducted using SAS 9.4 (SAS Institute, Cary, NC).

## Results:

Among the 1064 participants recruited into the LURN Observational Cohort Study, 920 had complete data on metabolic factors and had complete responses to at least 15 out of 22 questions on the LUTS Tool. Demographics and characteristics of these 920 participants (456 males and 464 females) are included in the Supplemental Table. The mean age was  $59.1 \pm 13.9$  years; 81.5% were white, most had central obesity per ATP III definition (60.4%) or IDF definition (70.2%), 43.4% had general obesity (BMI  $\geq 30$ ), 76.5% were overweight (BMI  $\geq 25$ ), 65.2% had hypertension, 31.5% had dyslipidemia, and 17.1% had diabetes mellitus. The lower percentage with central obesity using the ATP III versus the IDF definition was due to higher cutoff values with ATP III. Regarding incontinence, 33.1% had no UI, 7.4% had SUI, 20.4% had UUI, 26.1% had mixed UI, and 13.0% had other UI; 63.4% of participants had OAB (urgency plus either frequency or nocturia). Among those with OAB, 66.4% had OAB with UUI (“OAB-wet”), and 33.6% had OAB without UUI (“OAB-dry”).

Table 1 shows the relationships between central obesity and LUTS (multivariable logistic regression). We include the results from all three obesity measures in the table (central obesity cutoffs defined by ATP III, defined by IDF, and waist circumference as a continuous variable), but will describe the results of the continuous predictors in more detail below, keeping in mind that the dichotomous predictors also identify several significant predictors.

Higher waist circumference was associated with higher odds of having any UI (OR=1.16 per 10 cm increase,  $p=0.008$ ) and UUI (OR=1.24 per 10 cm increase,  $p=0.001$ ) in both sexes, and SUI in females (OR=1.27 per 10 cm increase,  $p=0.008$ ). In females, there was a relationship between waist circumference and OAB (OR=1.25/10 cm increase,  $p=0.003$ ),

however the relationship was not detected in males (OR=1.02/10 cm increase,  $p=0.73$ ). Among participants with OAB, higher waist circumference was associated with UI versus without UI (“OAB-wet” vs. “OAB-dry”, OR=1.23/10 cm increase,  $p=0.006$ ). Higher waist circumference was additionally associated with higher odds of frequency and nocturia 1 and 2 (OR=1.16/10cm,  $p=0.009$ ; OR=1.16,  $p=0.035$ ; OR=1.12,  $p=0.016$ , respectively). Dyslipidemia was also associated with nocturia 2 in this model (OR=1.46,  $p=0.035$ )

Table 2 shows the relationships between general obesity, overweight status, continuous BMI, and LUTS (multivariable logistic regression). As above with waist circumference, we include the results from all three obesity measures in the table (BMI  $\geq 30$ , BMI  $\geq 25$ , BMI as a continuous variable), but will describe the results of the continuous predictors in more detail below, keeping in mind that the dichotomous predictors also identify several significant predictors.

Higher BMI was associated with any UI (OR=1.20/5 unit increase,  $p=0.006$ ) and UI (OR=1.31/5 unit increase,  $p<0.001$ ) in both sexes, and SUI in females (OR=1.32/5 unit increase,  $p=0.009$ ). In females, there was a relationship between BMI and OAB (OR=1.38/5 unit increase,  $p<0.001$ ), however again no association was detected in males (OR=1.05/5 unit increase,  $p=0.59$ ). Among participants with OAB, higher BMI was associated with UI versus without UI (“OAB-wet” vs. “OAB-dry”, OR=1.26/5 unit increase,  $p=0.006$ ). Dyslipidemia was also associated with nocturia 2 in this model (OR=1.48,  $p=0.021$ ). Of the other covariates, older age, female sex, and African American race were also associated with multiple urinary symptoms (Tables 1 and 2).

## Discussion:

Previous community-based studies have demonstrated relationships between high BMI and UI in females,<sup>7</sup> and between metabolic factors and higher AUA Symptom Index in males.<sup>7</sup> However, the relationships between metabolic factors and OAB in females were less clear. Also the relationships between metabolic factors and UI and OAB in males have not been well studied. In this study we found associations between both central and general obesity and OAB in females, and UI and its subtypes in males and females. We did not find association between central or general obesity and OAB in males. We also found an association between dyslipidemia and nocturia 2 in the clinical population.

Our observations in the clinical cohort were consistent to those reported in the Boston Area Community Health (BACH) population-based cohort. In the BACH Study, the odds of having OAB increased with waist circumference and with BMI in females. However, in males, there were negative associations up to 100 cm waist circumference or a BMI of 27.5, and then positive associations beyond those thresholds. In longitudinal studies, Dallosso et al also showed that obesity at baseline was associated with new onset of OAB one year later in women; however obesity at baseline was not associated with new onset of OAB one year later in men. Taken together, the data suggested that there might be important sex difference in the relationships between obesity and OAB.

This sex difference raises some interesting research possibilities. For example, hormonal factors may play a role. Anatomic factors may account for some of the sex differences. Increased abdominal pressure from central (visceral) obesity could increase the pressure on the bladder, stretch the pelvic floor, or trigger urine entry into the proximal urethra, which could in turn cause prolapse, urethral hypermobility, and OAB symptoms in females. Males however are less susceptible to these anatomic forces since males do not develop prolapse, urethral hypermobility, or an open bladder neck. Besides local mechanical forces, systemic factors such as neuroendocrine, vascular or inflammatory processes can provide a link between obesity and OAB. Adipose tissue produces leptin, which can stimulate the sympathetic nervous system activity, and exacerbate urinary frequency. The generation of inflammatory factors (e.g. cytokines, C-reactive protein) from visceral adipose tissue may also influence bladder sensory function. Obesity is also associated with vascular dysfunction and ischemia. Vascular ischemia in pelvic structures including the bladder has been demonstrated to cause detrusor overactivity and OAB in animal models. In summary, the pathophysiology that links obesity and OAB is poorly understood, and the mechanisms behind the sex differences need to be further examined.

Among participants with OAB, increases in continuous BMI and waist circumference were associated with UII versus without UII (“OAB-wet” vs. “OAB-dry”). Our results were in contrast to a Korean community study which showed that high BMIs were associated with “OAB-dry” but not “OAB-wet”. Our clinical cohort consisted of patients who presented to tertiary urology and urogynecology clinics seeking care for their LUTS; thus, it was possible that the community sample may have less severe UII, and therefore did not permit the discovery of an association.

Although a number of measures of adiposity are available that account for different aspects of obesity, they have rarely been evaluated together for their associations with urinary symptoms. Among studies that have examined obesity, most have only reported on BMI. Since the distribution of fat in the abdominal viscera – central obesity – is a key component of metabolic syndrome, and central obesity is more predictive of cardiovascular risk than high BMI, we have specifically examined three measures of central obesity (the ATP III definition, the IDF definition, and waist circumference as a continuous measure) and three measures of BMI (general obesity [BMI  $\geq 30$ ], overweight status [BMI  $\geq 25$ ], and BMI as a continuous measure), to assess differences among these measures of obesity with respect to their relationships to UI and OAB. Overall the continuous measures of waist circumference or BMI showed the strongest relationships with incontinence measures, followed by central obesity (ATP III) and general obesity (BMI  $\geq 30$ ). As shown in Figure 1, there was high correlation between waist circumference and BMI in our care seeking cohort. No associations were found with central obesity defined by IDF, which has lower cutoffs than ATP III. This is likely due to the distribution of waist circumference in the LURN sample resulting in similar odds of LUTS when participants are grouped using the IDF definition despite a positive association with waist circumference (as a continuous variable). This is illustrated in Figure 2 using UII in males as an example. It is also possible that the positive association is less strong at lower values of waist circumference, however we did not have adequate sample size to detect this difference. Despite this limitation, the use of continuous predictor variables for waist circumference and BMI revealed the substantial gain in power

over use of dichotomized variables, and may explain some of the conflicting results from prior studies that used dichotomous predictors. For similar reasons, overweight status was not associated with many LUTS due to lower BMI cutoffs, even though there were positive associations with higher BMI (as a continuous variable).

In this study we were not able to demonstrate any relationships to other metabolic factors including diabetes and hypertension. A novel association between dyslipidemia and nocturia<sup>2</sup> was observed, that to our knowledge has never been reported in the literature. One study from Taiwan (n=1827) found an increased odds of OAB in individuals with dyslipidemia. In the BACH Study, the use of statins was associated with lower prevalence of LUTS among older men but not among women or younger men. Unfortunately neither of these studies had looked specifically at nocturia. The significance of the potential link between nocturia and dyslipidemia is unclear at this time and should be further investigated.

Our data showed that obesity is a key modifiable metabolic factor associated with UI, SUI, UUI, and OAB. This finding may have important implications for primary and secondary preventive strategies to reduce the prevalence and burden of OAB and UI. Weight loss interventions targeting physical activity and healthy diet might be considered as an adjuvant therapy in OAB and UI patients who are obese. A 2015 Cochrane systematic review concluded that the therapeutic effect of weight loss on UI is building and should be a research priority.

The strength of the study included: enrollment of a large cohort (>900) of patients who were seeking care for their LUTS at six clinical centers; inclusion of both males and females; and measurement of both waist circumference and BMI so we could examine various measures of obesity along with other metabolic factors. Our use of continuous predictor variables for waist circumference and BMI also revealed the substantial gain in power over use of dichotomized variables. There are several potential weaknesses of the study. We did not collect fasting glucose, hemoglobin A1c, HDL-cholesterol, or triglyceride serum levels to fully assess metabolic syndrome. For participants who took medications for hypertension, dyslipidemia or diabetes, we did not further sub-categorize them into those whose conditions were poorly controlled versus well controlled. Also, instead of using multiple instruments to assess different LUTS (e.g. OAB-q for OAB), we chose to use one instrument (LUTS Tool) as a comprehensive tool to assess a variety of LUTS. Only participants from specialty clinics (urology, urogynecology) in tertiary academic medical centers were enrolled, which may limit the generalizability of the findings. Finally our cross-sectional data does not permit one to draw inference on causality or temporal relationships.

## Conclusions:

In care-seeking patients with LUTS, central obesity and general obesity were key metabolic factor associated with UI in both males and females, and with OAB in females. We did not demonstrate association between obesity and OAB in males. The association between dyslipidemia and nocturia<sup>2</sup> needs further research. Weight loss is a promising therapeutic intervention. Future research could focus on optimizing this intervention.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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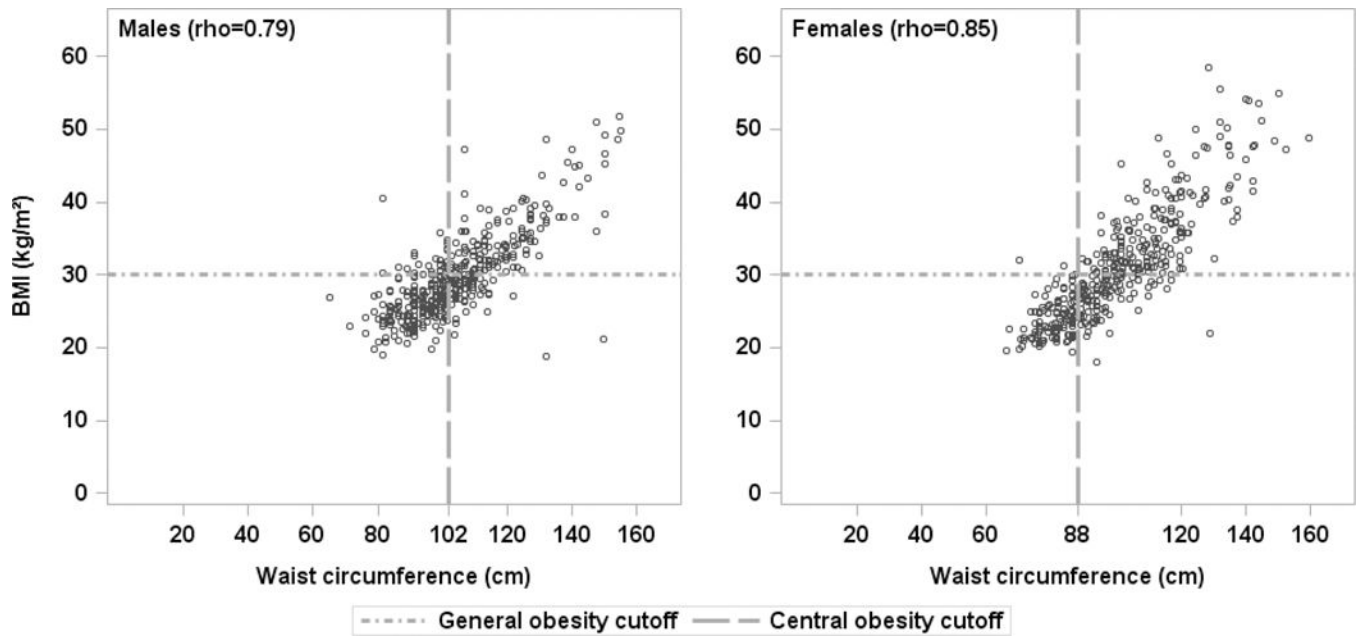
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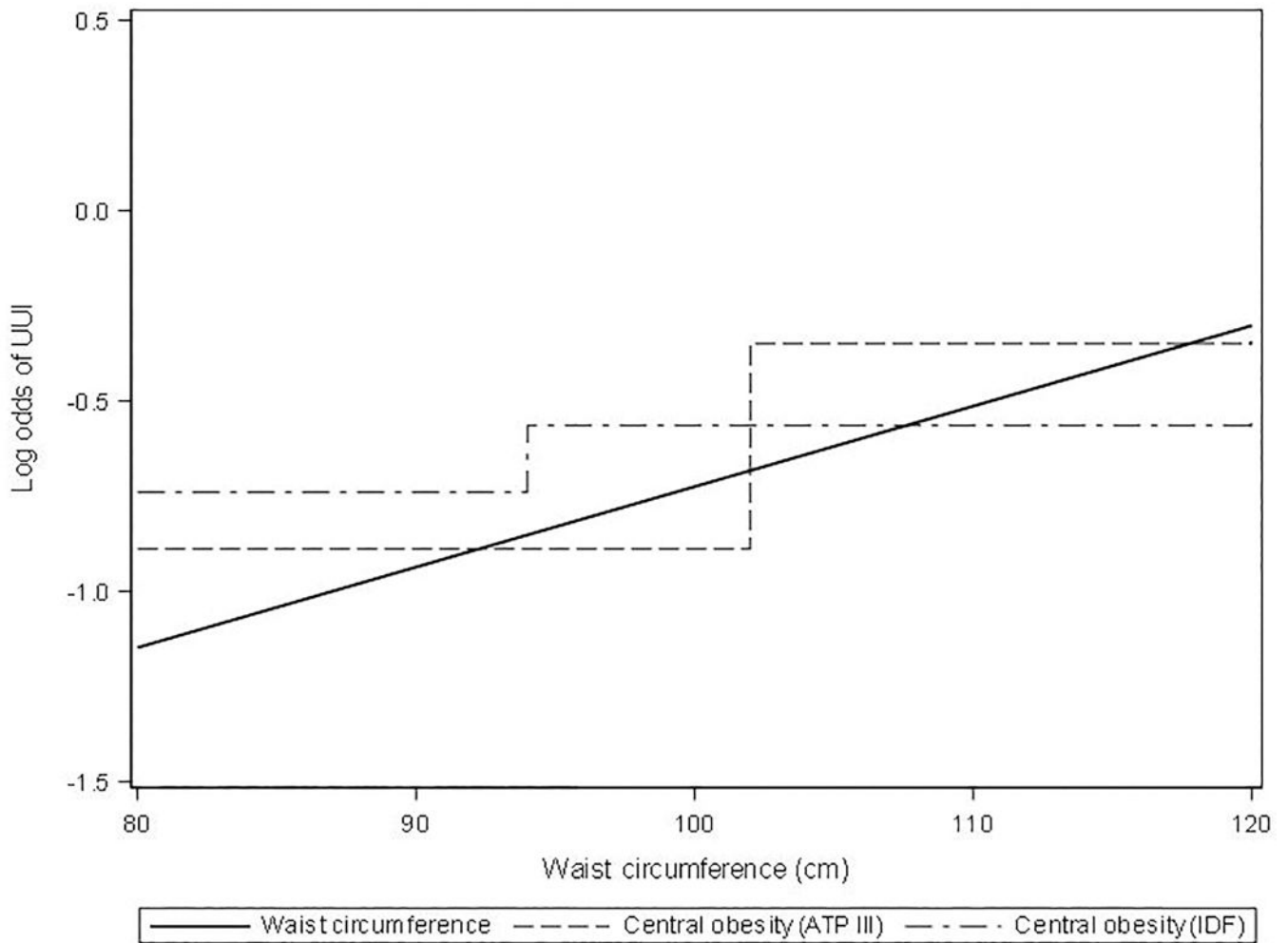
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**Figure 1: Scatterplots of body mass index (BMI) by waist circumference for males and females with Pearson correlation coefficients (both  $p < 0.001$ ). Dashed lines show the ATP III criteria for central obesity (waist circumference 102 cm for males, 88 cm for females) and the general obesity criterion of BMI 30.**



**Figure 2: Odds ratio of UUI in males for various measures of central obesity.**

When waist circumference is included as a continuous variable, the log odds of UUI increases as waist circumference increases (solid line). When waist circumference is dichotomized, the averaged log odds is different enough that the difference is significant with the ATP III cutoffs (dashed line), but not with the IDF cutoffs (dashed-dotted line).

**Table 1:** Relationships between central obesity (waist circumference) and LUTS (multivariable logistic regression)

Dependent Variable	Models using Central Obesity ATP III definition (Yes vs. No)					Models using Central Obesity IDF definition (Yes vs. No)					Models using Waist Circumference (continuous measures)				
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
UI (odds of having any UI)	Central Obesity	1.411	1.009	1.973	0.058	Central Obesity	0.872	0.585	1.300	0.637	Waist Circumference (per 10 cm)	1.155	1.048	1.273	0.008
	Female vs. Male	4.572	3.223	6.487	<0.001	Female vs. Male	0.207	0.146	0.293	<0.001	Female vs. Male	5.262	3.733	7.418	<0.001
SUI (in Females only) #	Central Obesity	2.026	1.102	3.725	0.036	Central Obesity	1.340	0.541	3.318	0.642	Waist Circumference (per 10 cm)	1.271	1.08	1.495	0.008
	Central Obesity	1.716	1.175	2.507	0.011	Central Obesity	1.087	0.689	1.713	0.807	Waist Circumference (per 10 cm)	1.236	1.105	1.382	0.001
	Female vs. Male	6.207	4.247	9.073	<0.001	Female vs. Male	7.174	4.868	10.572	<0.001	Female vs. Male	7.761	5.302	11.359	<0.001
UUI	Black vs. Non-Black	2.157	1.174	3.963	0.026	Hypertension (Yes vs. No)	1.548	1.039	2.305	0.059	Black vs. non-Black	2.144	1.166	3.94	0.022
	Central Obesity – Males	1.023	0.684	1.530	0.911	Central Obesity	1.232	0.837	1.814	0.407	Waist Circumference (per 10 cm) – Males	1.022	0.902	1.159	0.731
OAB	Central Obesity – Females	2.493	1.530	4.063	0.001	Female vs. Male	2.123	1.546	2.914	<0.001	Waist Circumference (per 10 cm) – Females	1.248	1.093	1.426	0.003
	Age (per 10 years)	1.174	1.050	1.312	0.011	Age (per 10 years)	1.182	1.057	1.321	0.011	Age (per 10 years)	1.162	1.042	1.295	0.013
	Black vs. Non-Black	1.919	1.117	3.298	0.031	Black vs. Non-Black	1.947	1.137	3.333	0.033	Black vs. non Black	1.881	1.098	3.221	0.031

Dependent Variable	Models using Central Obesity ATP III definition (Yes vs. No)					Models using Central Obesity IDF definition (Yes vs. No)					Models using Waist Circumference (continuous measures)				
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
<b>OAB with UII vs. without UII (among those with OAB only) <sup>A</sup></b>	Central Obesity	1.422	0.931	2.172	0.123	Central Obesity	1.159	0.685	1.96	0.681	Waist Circumference (per 10 cm)	1.225	1.076	1.395	0.006
	Female vs. Male	4.063	2.681	6.159	<0.001	Female vs. Male	4.378	2.908	6.591	<0.001	Female vs. Male	5.016	3.345	7.521	<0.001
<b>Urgency Symptoms</b>	Central Obesity	1.327	0.96	1.834	0.108	Central Obesity	1.036	0.698	1.536	0.862	Waist Circumference (per 10 cm)	1.086	0.986	1.195	0.096
	Female vs. Male	2.791	2.008	3.88	<0.001	Female vs. Male	3.122	2.237	4.356	<0.001	Female vs. Male	3.144	2.273	4.347	<0.001
	Black vs. Non-Black	1.784	1.018	3.126	0.058	Black vs. Non-Black	1.714	0.978	3.003	0.098	Black vs. non-Black	1.695	0.968	2.968	0.073
	Hypertension (Yes vs. No)	1.548	1.089	2.201	0.027	Hypertension (Yes vs. No)	1.464	1.049	2.043	0.050	Hypertension (Yes vs. No)	1.417	1.01	1.986	0.051
<b>Frequency Symptoms</b>	Central Obesity	1.223	0.862	1.736	0.282	Central Obesity	1.042	0.678	1.602	0.862	Waist Circumference (per 10 cm)	1.157	1.046	1.28	0.009
	Black vs. Non-Black	2.111	1.063	4.191	0.048	Black vs. non-Black	2.103	1.059	4.177	0.059	Black vs. non-Black	2.142	1.08	4.247	0.036
	Hypertension (Yes vs. No)	1.548	1.089	2.201	0.027	Hypertension (Yes vs. No)	1.593	1.122	2.261	0.026					
<b>Nocturia <sup>1</sup></b>	Central Obesity – Males	1.138	0.533	2.430	0.769	Central Obesity	1.094	0.607	1.970	0.825	Waist Circumference (per 10 cm)	1.161	1.018	1.322	0.035
	Central Obesity – Females	2.787	1.563	4.969	0.001	Female vs. Male	0.528	0.322	0.865	0.026	Female vs. Male	0.558	0.351	0.889	0.022
	Age (per 10 years)	1.367	1.171	1.597	<0.001	Age (per 10 years)	1.334	1.136	1.567	0.002	Age (per 10 years)	1.377	1.184	1.601	<0.001
	Dyslipidemia (Yes vs. No)					Dyslipidemia (Yes vs. No)	1.714	0.927	3.17	0.127					

Dependent Variable	Models using Central Obesity ATP III definition (Yes vs. No)					Models using Central Obesity IDF definition (Yes vs. No)					Models using Waist Circumference (continuous measures)				
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
Nocturia	Central Obesity	1.201	0.890	1.620	0.263	Central Obesity	1.203	0.813	1.781	0.473	Waist Circumference (per 10 cm)	1.122	1.029	1.225	0.016
	Age (per 10 years)	1.374	1.226	1.54	<0.001	Female vs. Male	0.762	0.560	1.038	0.127	Female vs. Male	0.775	0.578	1.04	0.096
	Black vs. Non-Black	2.883	1.703	4.882	<0.001	Age (per 10 years)	1.352	1.204	1.517	<0.001	Age (per 10 years)	1.336	1.194	1.495	<0.001
	Dyslipidemia (Yes vs. No)	1.605	1.153	2.232	0.011	Black vs. non-Black	2.873	1.695	4.87	<0.001	Black vs. non-Black	2.739	1.616	4.643	0.001
						Dyslipidemia (Yes vs. No)	1.545	1.105	2.159	0.026	Dyslipidemia (Yes vs. No)	1.455	1.044	2.026	0.035

# Males with a history of prostate cancer were excluded from the study. Since male SUI was rare, only female SUI was analyzed.

<sup>^</sup> Comparing OAB with UUI versus OAB without UUI (“OAB-wet” versus “OAB-dry”).

Abbreviations: CI = confidence limit; FDR = false discovery rate; OAB = overactive bladder syndrome; SUI = stress urinary incontinence; UI = urinary incontinence; UUI = urgency urinary incontinence

**Table 2:** Relationships between general obesity (BMI), overweight status and LUTS (multivariable logistic regression)

Dependent Variable	Models using General Obesity (BMI 30 vs. <30)					Models using Overweight Status (BMI 25 vs. <25)					Models using BMI (continuous measures)				
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
UI (odds of having any UI)	General Obesity	1.443	1.058	1.968	0.028	Overweight	1.323	0.922	1.898	0.160	BMI (per 5 units)	1.202	1.063	1.359	0.006
	Female vs. Male	4.644	3.386	6.369	<0.001	Female vs. Male	4.832	3.518	6.636	<0.001	Female vs. Male	4.641	3.382	6.369	<0.001
SUI (in Females only) #	General Obesity	1.931	1.132	3.296	0.023	Overweight	1.395	0.792	2.457	0.283	BMI (per 5 units)	1.318	1.087	1.599	0.009
	General Obesity	1.857	1.316	2.619	0.001	Overweight	1.280	0.840	1.949	0.283	BMI (per 5 units)	1.305	1.142	1.492	<0.001
UII	Female vs. Male	6.387	4.526	9.012	<0.001	Female vs. Male	7.081	4.973	10.084	<0.001	Female vs. Male	6.377	4.514	9.009	<0.001
	General Obesity	1.157	0.787	1.699	0.502	Hypertension (Yes vs. No)	1.487	1.015	2.178	0.059	Female vs. Male	1.047	0.886	1.238	0.588
OAB	General Obesity – Males	2.432	1.576	3.753	<0.001	Overweight	1.778	1.285	2.461	0.001	BMI (per 5 units) Male	1.385	1.182	1.623	<0.001
	General Obesity – Females	1.141	1.030	1.263	0.019	Female vs. Male	2.220	1.665	2.960	<0.001	BMI (per 5 units) Female	1.135	1.025	1.257	0.021
OAB with UII vs. those with OAB only) <sup>^</sup>	Age (per 10 years)	1.789	1.22	2.624	0.007	Age (per 10 years)	1.129	1.019	1.251	0.037	Age (per 10 years)	1.26	1.079	1.472	0.006
	General Obesity	1.789	1.22	2.624	0.007	Overweight	1.132	0.708	1.811	0.628	BMI (per 5 units)	1.26	1.079	1.472	0.006
OAB with UII vs. those with OAB only) <sup>^</sup>	General Obesity	4.498	3.073	6.585	<0.001	Female vs. Male	4.716	3.230	6.885	<0.001	Female vs. Male	4.449	3.039	6.512	<0.001
	Female vs. Male	4.498	3.073	6.585	<0.001	Female vs. Male	4.716	3.230	6.885	<0.001	Female vs. Male	4.449	3.039	6.512	<0.001



Dependent Variable	Models using General Obesity (BMI 30 vs. <30)					Models using Overweight Status (BMI 25 vs. <25)					Models using BMI (continuous measures)				
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
<b>Urgency Symptoms</b>	General Obesity – Males	1.217	0.828	1.789	0.366	Overweight	1.671	1.190	2.346	0.008	BMI (per 5 units)	1.22	1.083	1.375	0.003
	General Obesity – Females	2.573	1.566	4.228	0.001	Female vs. Male	3.009	2.226	4.066	<0.001	Female vs. Male	2.812	2.086	3.792	<0.001
	Black vs. Non-Black	1.742	1.022	2.969	0.050	Black vs. Non-Black	1.790	1.052	3.046	0.052					
		1.597	1.149	2.22	0.010	Overweight	1.229	0.848	1.782	0.300	BMI (per 5 units)	1.227	1.077	1.397	0.005
<b>Frequency Symptoms</b>	Black vs. Non-Black	2.248	1.172	4.313	0.023	Black vs. Non-Black	2.266	1.181	4.350	0.028	Black vs. non-Black	2.19	1.14	4.207	0.026
						Hypertension (Yes vs. No)	1.405	1.004	1.968	0.062					
	General Obesity	1.132	0.737	1.74	0.598	Overweight	1.600	1.015	2.523	0.059	BMI (per 5 units) Males	0.792	0.598	1.049	0.116
<b>Nocturia 1</b>	Female vs. Male	0.525	0.335	0.823	0.010	Female vs. Male	0.544	0.347	0.853	0.019	BMI (per 5 units) Female	1.185	0.986	1.425	0.082
	Age (per 10 years)	1.311	1.131	1.521	0.001	Age (per 10 years)	1.305	1.124	1.515	0.001	Age (per 10 years)	1.316	1.132	1.529	0.001
	Dyslipidemia (Yes vs. No)	1.894	1.057	3.394	0.041	Dyslipidemia (Yes vs. No)	1.832	1.020	3.289	0.059	Dyslipidemia (Yes vs. No)	1.854	1.031	3.335	0.047
	General Obesity	1.028	0.779	1.357	0.845	Overweight	0.993	0.716	1.377	0.966	BMI (per 5 units)	1.029	0.93	1.139	0.588
<b>Nocturia 2</b>	Age (per 10 years)	1.374	1.236	1.528	<0.001	Age (per 10 years)	1.355	1.218	1.507	<0.001	Age (per 10 years)	1.355	1.218	1.507	<0.001
	Black vs. Non-Black	2.833	1.728	4.643	<0.001	Black vs. Non-Black	2.862	1.741	4.703	<0.001	Black vs. non-Black	2.815	1.713	4.626	<0.001
						Female vs. Male	0.728	0.551	0.961	0.044	Female vs. Male	0.723	0.547	0.955	0.03

Dependent Variable	Models using General Obesity (BMI <30 vs. <30)				Models using Overweight Status (BMI <25 vs. <25)				Models using BMI (continuous measures)						
	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value	Variable	Odds Ratio (OR)	Lower 95% CI	Upper 95% CI	FDR adjusted p-value
Dyslipidemia (Yes vs. No)		1.547	1.135	2.109	0.010	Dyslipidemia (Yes vs. No)	1.493	1.092	2.039	0.026	Dyslipidemia (Yes vs. No)	1.478	1.081	2.021	0.021

# Males with a history of prostate cancer were excluded from the study. Since male SUI was rare, only female SUI was analyzed.

^ Comparing OAB with UUI versus OAB without UUI (“OAB-wet” versus “OAB-dry”).

Abbreviations: CI = confidence limit; FDR = false discovery rate; OAB = overactive bladder syndrome; SUI = stress urinary incontinence; UI = urinary incontinence; UUI = urgency urinary incontinence