Postmenozal Kadınlarda Visseral Adipozite İndeksi ve Aşırı Aktif Mesane: Yeni Bir Risk Belirteci

Visceral Adiposity Index and Overactive Bladder in Postmenopausal Woman: A Novel Predictive Risk Factor

Hüseyin Eren1, Mustafa Ozan Horsanlı2, Eyüp Dil1, Emin Özbek3

1Recep Tayyip Erdoğan Üniversitesi Tıp Fakültesi Üroloji Anabilim Dalı, Rize, Türkiye
2Recep Tayyip Erdoğan Üniversitesi Eğitim Ve Araştırma Hastanesi Üroloji Kliniği, Rize, Türkiye
3İstanbul Üniversitesi Cerrahpaşa Tıp Fakültesi, İstanbul, Türkiye

ÖZ
AMAC: 45 yaş üstü postmenopozal kadın hastalarda visseral adipozite indeksi (VAI) ile aşırı aktif mesane semptomları arasındaki ilişkiyi araştırmak.


BULGULAR: Hastaların yaş ortalaması 57.2 ± 8.3 yıl, ortalama vücut kitle indeksi (VKİ) 32.3 ± 2.8 kg/m2 idi. Gruplar arasında Pabd, post miksiyonel rezidü idrar miktarı (PVR), maksimum mesane kapasitesi (MMK) ve OAB-8 skoru açısından anlamlı fark bulundu. VAI ile PVR, detrusor aşırı aktivitesi, idrar sıklığı, urge inkontinans, noktülü incontinans ve OAB-8 skoru arasındaki ilişki yönden negatif korelasyon tesbit edildi.

SONUC: VAI düzeyi yüksek olan hastalarda daha fazla aşırı aktif mesane semptomu olduğu ve daha kötü urodinamik bulguları olduğunu gözlemledik. VAI ayrıca aşırı aktif mesane semptomlarının önünümlerine bir belirteç olarak kullanılabilir.

Anahtar kelimeler: Obezite; Aşırı aktif mesane; Üriner inkontinans; Visceral Adipozite İndeksi; Bel çevresi; Detrusör aşırı aktivitesi

ABSTRACT
INTRODUCTION: To investigate the association between visceral adiposity index (VAI) and overactive bladder (OAB) symptoms in postmenopausal female patients aged over 45 years.

MATERIALS and METHODS: Between March 2017 and December 2017, 76 postmenopausal female patient with OAB symptoms were evaluated. Anthropometric indices serum cholesterol levels, fasting glucose levels, urodynamics findings and OAB-8 scores were recorded. VAI was calculated according to gender-specific formula. Participants were divided into three groups according to VAI levels. The relationship between VAI and OAB symptoms were statistically compared in terms of urodynamical outcomes and OAB-8 scores.

RESULTS: Mean age of the patients were 57.2 ± 8.3 years and mean BMI was 32.3 ± 2.8 kg/m2. Statistical significance was observed in terms of abdominal pressure (Pabd), post voiding residual urine mounts (PVR), maximum bladder capacity (MBC) and OAB-8 scores between groups. VAI correlated positively with PVR, detrusor overactivity (DOA), urinary frequency, urge incontinence, nocturia and OAB-8 scores whereas a negative correlation existed MBC and first desire to void.

CONCLUSIONS: We observed that patients with higher levels of VAI had more OAB symptoms and worse urodynamical findings. VAI may also be used as a marker for predicting overactive bladder symptoms.

Keywords: Obesity; Overactive bladder; Urinary Incontinence; Visceral Adiposity Index; Waist Circumference; Detrusor overactivity
INTRODUCTION

According to the International Continence Society (ICS), overactive bladder (OAB) is determined as a urinary urgency, usually accompanied by increased daytime frequency and nocturia, with urinary incontinence or without, in the absence of urinary tract infection or other detectable disease causing distress and negative impact on a person’s quality of life (1). International population-based studies have reported prevalence rates of OAB approximately 12% (6,7) and it rises 17% in patients aged 40 and over (2). OAB symptoms are not only disturbs patients but also have a significant negative impact on many aspects of the lives of individuals who represent a particularly effective health burden in terms of quality of life and productivity (3). It is known that obesity is a strong risk factor for OAB. Several population-based studies found that increased body mass index (BMI) could trigger symptoms of OAB (4, 5).

The increasing prevalence of obesity and the obesity-related complications are health problems growing in developed countries (6). Although obesity was often described as BMI, previous studies showed that abdominal fat distribution might be a more prominent marker of obesity-related pathological features (7). The adipose tissue is mainly deposited into subcutaneous and visceral compartments. Visceral fat tissue consists mainly from omental fat and mesenteric fat. They have a higher metabolic activity than subcutaneous adipose tissue (8). Visceral adipose tissue has endocrine, metabolic, and immunological functions. These endocrine, metabolic and immunologic functions cause pathogenesis of the pro-inflammatory, procoagulant state of metabolic syndrome associated with insulin resistance (8). Visceral adipose index (VAI) is a new mathematical model that predicts visceral adiposity based on anthropometric and lipid profiles. It is a reliable formula of cardiometabolic risk which is useful indicator of visceral fat function without using costly and inappropriate imaging studies (9). In the present study, we aimed to evaluate the relationship between VAI and OAB symptoms with or without urinary incontinence.

MATERIALS and METHODS

In the present study, we evaluate patient charts of postmenopausal patients who had urodynamic study by clinician’s preferences with OAB symptoms (n=76), retrospectively. The collected data was recorded between March to December 2017. All patients were over 45-years-old. Female patients with a complete anthropometric indices, incontinence, urodynamic study datas and laboratory results including serum cholesterol, fasting glucose were included study. Detailed medical histories of all patients were obtained before the study. Height, weight, and waist circumference (WC) were measured. BMI was computed by the formula as the ratio of weight to the square of height (kg/m2). WC was measured on the line between the iliac crest and the lower costal margin parallel to the ground after subjects exhaled.

Each patient filled out OAB questionnaire-8 Turkish validated short form and all datas were recorded. Inclusion criteria were age over 45, postmenopausal female gender, OAB symptoms with or without urinary incontinence, and available urodynamic study. Exclusion criteria were pregnancy, alcohol intake, bladder cancer history, previous incontinence surgery, presence of neurological disease, urinary infection, bladder stone, pelvic radiotherapy history and taking of estrogen therapy (n=54). VAI was calculated according to gender-specific formula in figure-1(9).

Figure 1. Visceral adiposity index formula for female patients

Females: $\text{VAI} = \left( \frac{\text{WC}}{36.58 + (1.89 \times \text{BMI})} \right) \times \left( \frac{\text{TG}}{0.81} \right) \times \left( \frac{1.52}{\text{HDL}} \right)$

VAI: Visceral adiposity index; WC: Waist circumference; BMI: Body mass index; TG: Triglyceride; HDL: High-density lipoprotein

Patients were divided into three groups according to VAI levels such as standard 1-9 (group 1), slightly high 10-14 (group 2), and high 15-30 (group 3) (10).

Urodynamic studies were performed according to good urodynamics principles. The urodynamic study
consists of free flow uroflowmetry, filling cystometry in semi-sitting position, stress urine leak test and pressure-flow study. The bladder was filled with saline at room temperature at a filling rate of 30 ml/min to prevent the detrusor overactivity due to filling rate. Peri-anal surface electromyography activity was monitored in both the filling and pressure flow phases of the urodynamic study. Abdominal pressure, post voiding residual urine (PVR), bladder capacity, detrusor over activity (DOA) and first desire to void parameters was recorded.

All statistical analyses were conducted by SPSS 17.0 statistical package program. Categorical variables were demonstrated as frequencies and percentages, continuous variables were demonstrated by mean±standard deviation. One way- ANOVA test was used to compare groups and post hoc Tukey test was used to examine the differences between groups. Pearson correlation analysis was used to test correlation between VAI and groups. A p value of < 0.05 was set for a statistical significance level.

## RESULTS

Seventy-six postmenopausal female patient aged over 45-years were included in the study. Mean age of the patients were 57.2±8.3 years and mean BMI was 32.3±2.8 kg/m2. Antropometric incides and laboratory characteristics of patients were summarized in (Table 1).

Thirty two (42.1%) patients had standard, 24 (31.6%) patients slightly high and 20 (26.3%) patients had high VAI levels. Although 6 (7.9%) of all patients were smoker, 70 (92.1%) were non-smoker. Thirty-three (43.4%) patients have frequency, 19 (25%) patients have urge urinary incontinence and 13 (17.1%) patients have nocturnal symptoms. Urodynamic study results for Pabd (group 1 to 2), PVR (group 1 to 3), and maximum bladder capacity (MBC) (group 1 to 2) parameters were statistically significantly different between groups (Table 2).

A urodynamic DOA rate was 11.8% (n=9/76) in the in the whole group. DAO recorded in 3.1% (n=1), 12.5% (n=3), and 25% (n=5) of patients for Group 1, Group 2 and Group 3, respectively. Recorded urodynamic DOA and OAB-8 scores were statistically significantly higher in group 3 than group 1 and 2 (Table 2). Positive correlation was found between groups in terms of PVR, DOA, urinary frequency, urge incontinence, nocturia and OAB-8 scores parameters in Pearson correlation test, and negative correlation was found in MBC and first desire to void in urodynamic study (Table 3).

### DISCUSSION

The underlying pathophysiology of OAB is still unclear, but it may be due to abnormalities in the detrusor, spontaneous motor activity abnormalities, triggered urgency and neurotransmitter release from urothelium by central mechanisms, especially cerebral white matter changes in prefrontal regions of the brain. The urothelium-suburothelium compartment of the bladder has a high metabolic activity and it is considered to responsible with mechanosensory transduction (11). But, the underlying pathology and relationship between obesity and OAB is not clear and it is still controversial.
Table 2. Comparison of groups according to urodynamic study features and Overactive Bladder-8 Turkish validated questionnaire scores

<table>
<thead>
<tr>
<th></th>
<th>Group 1</th>
<th>Group 2</th>
<th>Group 3</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Pdet, (cm H2O), mean± SD</strong></td>
<td>18.05±13.7</td>
<td>18.9±12.6</td>
<td>16.2±15.5</td>
</tr>
<tr>
<td><strong>Pves, (cm H2O), mean± SD</strong></td>
<td>19.8±14.5</td>
<td>27.8±19.6</td>
<td>24.8±13.5</td>
</tr>
<tr>
<td><strong>Pabd, (cm H2O), mean± SD</strong></td>
<td>7.2±8.5b</td>
<td>15.1±15.1</td>
<td>8.8±3.8</td>
</tr>
<tr>
<td><strong>DOA, n (%)</strong></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Absent</td>
<td>31(96.9%)</td>
<td>21(87.5%)</td>
<td>15(75%)</td>
</tr>
<tr>
<td>Present</td>
<td>1(3.1%)</td>
<td>3 (12.5%)</td>
<td>5(25%)</td>
</tr>
<tr>
<td><strong>PVR (ml), mean± SD</strong></td>
<td>60.1±66.7a</td>
<td>85.4±90.6</td>
<td>153.5±139.3</td>
</tr>
<tr>
<td><strong>ALPP (mmHg), mean± SD</strong></td>
<td>28.6±28.8</td>
<td>31.8±17.5</td>
<td>27.9±26.6</td>
</tr>
<tr>
<td><strong>VLPP (mmHg), mean± SD</strong></td>
<td>30.3±27.8</td>
<td>38.2±21.5</td>
<td>34.4±27.08</td>
</tr>
<tr>
<td><strong>MBC (ml), mean± SD</strong></td>
<td>505.6±207.3b</td>
<td>444.4±202.4</td>
<td>351±168.5</td>
</tr>
<tr>
<td><strong>OAB-8 Score, mean± SD</strong></td>
<td>11.03±6.02b</td>
<td>14.7±7.2</td>
<td>19.05±9.9</td>
</tr>
</tbody>
</table>

**DOA**: Detrussor overactivity; **PVR**: Post Voiding residual urine; **ALPP**: Abdominal Leak Point Pressure; **VLPP**: Valsalva Leak Point Pressure; **MBC**: Maximum Bladder Capacity

*Significantly different between group 1 and 3, bSignificantly different between grades 1 and 2. p< 0.05 is considered statistically significant

Table -3 Correlations between study parameters

<table>
<thead>
<tr>
<th></th>
<th>VAI</th>
<th>MBC</th>
<th>PVR</th>
<th>DOA</th>
<th>OAB-8 score</th>
<th>FDV</th>
<th>UF</th>
<th>UUI</th>
<th>Nocturia</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>VAI</strong></td>
<td>1</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>MBC</strong></td>
<td></td>
<td>r:0.140</td>
<td>1</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>PVR</strong></td>
<td></td>
<td>r:0.404**</td>
<td>r:0.291*</td>
<td>1</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>DOA</strong></td>
<td></td>
<td>r:0.346**</td>
<td>r:0.275*</td>
<td>r:0.072</td>
<td>1</td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>OAB-8 score</strong></td>
<td></td>
<td>r:0.477**</td>
<td>r:0.104</td>
<td>r:0.399**</td>
<td>r:0.045</td>
<td>1</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>FDV</strong></td>
<td></td>
<td>r:0.235**</td>
<td>r:0.452**</td>
<td>r:0.055</td>
<td>r:0.217</td>
<td>r:0.220</td>
<td>1</td>
<td></td>
<td></td>
</tr>
<tr>
<td><strong>UF</strong></td>
<td></td>
<td>r:0.379**</td>
<td>r:0.110</td>
<td>r:0.200</td>
<td>r:0.132</td>
<td>r:0.069</td>
<td>r:0.227*</td>
<td>1</td>
<td></td>
</tr>
<tr>
<td><strong>UUI</strong></td>
<td></td>
<td>r:0.372**</td>
<td>r:0.029</td>
<td>r:0.348**</td>
<td>r:0.001</td>
<td>r:0.792**</td>
<td>r:0.219</td>
<td>r:0.659**</td>
<td>1</td>
</tr>
<tr>
<td><strong>Nocturia</strong></td>
<td></td>
<td>r:0.298**</td>
<td>r:0.073</td>
<td>r:0.316**</td>
<td>r:0.072</td>
<td>r:0.722**</td>
<td>r:0.099</td>
<td>r:0.519**</td>
<td>r:0.787**</td>
</tr>
</tbody>
</table>

**VAI**: Visceral adiposity index; **MBC**: Maximum bladder capacity; **PVR**: Post voiding residual urine; **DOA**: Detrussor over activity; **OAB**: Overactive bladder; **FDV**: First desire to void; **UF**: Urinary frequency; **UUI**: Urge urinary incontinence,

*p<0.05 is considered statistically significant; **p<0.01 is considered statistically significant

Visceral adiposity index is used as one of the indicator of visceral obesity. In the present study high visceral adiposity index levels were associated with OAB symptoms. Patients with higher VAI levels have increased PVR, frequency and urge urinary incontinence than patients with lower VAI levels. Urodynamic DOA rate was increased with increasing VAI. Although there are several studies reporting association between obesity and OAB, to our best of knowledge this is the first study evaluating relationship between VAI as a marker of central obesity and postmenopausal female patients with OAB.

BMI over 30 reported as an independent risk factor for OAB and DOA in some studies (13-16). In a study relationship between central and general obesity in terms of OAB and incontinence evaluated.
In this study central and general obesity cause urgency and urge incontinence in both gender, though over active bladder syndrome symptoms noted only for female gender (12). VAI is a reliable marker of central obesity. Our results supports findings of previously reported studies (13). Patients with high VAI levels have increases OAB symptoms, in terms of urgency and urge incontinence. In the present study we revealed that VAI could be used as a predictive anthropometric marker for OAB for postmenopausal females.

The relation between excessive weight gain and urinary incontinence were reported in some studies. All these studies concluded that excessive weight gain may increase the pressure on the pelvic floor muscles. This may lead to weakness of the urethral sphincter structure and cause urinary incontinence (14, 15). Excessive visceral fat tissue may cause bladder inflammation, possibly due to changes in neuroendocrine processes that can occur as urinary frequency and urgency. This suggests that the increase in visceral fat tissue may be a risk factor for stress urinary incontinence (4, 16). In a study with 8355 participants, overweight women found an increased risk of urinary incontinence compared to their weak counterparts. The risk increased 3% for per kilogram weight gain and 7% for each kilogram per meters2 for BMI. (17). In the present study, we support previous studies with positive correlation between VAI, urgency and urge incontinence.

In a study, 76 patients with LUTS were evaluated for pre and postoperative bariatric surgery. A statistically significant decrease in symptoms such as nocturia, urgency, stress urinary incontinence, frequency, unaware incontinence and, an increased quality of life scores were reported for women with LUTS at 6th week and 1st year after bariatric surgery (21). In the present study the mean BMI was 32.3±2.8 kg/m2. Patients with high BMI had higher symptoms scores and impaired urodynamic features. We consider that central obesity may have a role in underlying pathophysiology of OAB. As a result VAI may be a useful a marker for central obesity instead of BMI.

In a randomized control trial, 40 women with normal urodynamic results at baseline were examined by administering controlled weight loss as incontinence therapy. A statistically significant relation was reported with weight loss. Urodynamic findings were improved including decreased initial intravesical pressure, decreased intravesical pressure at maximum capacity and increased valsalva leak point pressure. For reason, present study also concludes that modification the risk factor changes the biological variant and intervenes in a reasonable mechanism (18). In present study we found similar results in terms of urodynamic findings. Patients with higher VAI levels, as an indicator of central obesity, have higher OAB-8 questionnaire scores, frequency, urge incontinence and nocturia. And also worse urodynamic findings such as increased PVR, DOA and first desire to void at low bladder filling.

Retrospective design and low number of patients were the most important limitations in the present study. We could also not explained the underlying pathophysiologcal mechanism between VAI and OAB. Histopathological changes on bladder wall or mucosa related to central obesity could not reveal. Although obesity is well known and modifiable risk factor for OAB in the literature, obesity related pathophysiological mechanism is still not clear.

**CONCLUSION**

We observed that patients with higher levels of VAI had more OAB symptoms and worse urodynamic findings. Therefore, VAI may be use as an indicator of central obesity and predictive marker for overactive bladder and follow-up VAI levels may be useful for the management of patients with overactive bladder.

**Conflict of interest**

Author declared no conflicts of interest

**REFERENCES**


