

Voiding Dysfunction

Effect of Obesity on Prostate-Specific Antigen, Prostate Volume, and International Prostate Symptom Score in Patients with Benign Prostatic Hyperplasia

Jong Min Kim, Phil Hyun Song, Hyun Tae Kim, Ki Hak Moon

Department of Urology, College of Medicine, Yeungnam University, Daegu, Korea

Purpose: We examined the correlation between body mass index (BMI) as a marker of obesity and prostate-specific antigen (PSA), prostate volume (PV), and International Prostate Symptom Score (IPSS) in patients with benign prostatic hyperplasia (BPH). **Materials and Methods:** From January 2008 to December 2008, we examined 258 patients diagnosed with BPH. Patients taking 5 α -reductase inhibitors or those diagnosed with prostate cancer were excluded from this study. BPH was defined as PV \geq 25 ml and IPSS \geq 8. BMI (kg/m²) was categorized into 4 groups as follows: BMI < 18.5 (underweight), BMI 18.5-23.0 (normal), BMI 23.0-27.5 (overweight), and BMI \geq 27.5 (obese). The relationships between PSA, PV, IPSS, and BMI were analyzed by correlation analysis and one-way ANOVA.

Results: The mean age of the patients was 65.19 ± 9.13 years and their mean BMI was 23.7 ± 4.4 kg/m². The mean PSA values of each BMI group were as follows: 3.42 ± 1.53 , 3.07 ± 1.88 , 2.74 ± 1.75 , and 2.60 ± 1.44 ng/ml. The PSA value was lowest in the obese group. The correlation analysis showed a negative correlation between BMI and PSA (Pearson's correlation coefficient=-0.142, p=0.023) and positive correlations between BMI and PSA is and PV (Pearson's correlation coefficient=0.32, p=0.001) and IPSS (Pearson's correlation coefficient=0.470, p=0.02). These correlations were also confirmed by one-way ANOVA.

Conclusions: Patients with an elevated BMI tended to have lower PSA values, larger PVs, and a higher IPSS. We suggest that weight loss could be helpful for BPH symptom relief as well as for detection of coexisting prostate cancer in BPH patients.

Key Words: Analysis of variance; Body mass index; Prostatic hyperplasia

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (http://creativecommons.org/licenses/by-nc/3.0) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

Article History: received 18 March, 2011 accepted 31 May, 2011

Corresponding Author:

Ki Hak Moon Department of Urology, Yeungnam University College of Medicine, 317–1, Daemyung 5-dong, Nam-gu, Daegu 705–717, Korea TEL: +82–53–620–3692 FAX: +82–53–627–5535 E-mail: khmoon@med.yu,ac.kr

INTRODUCTION

Benign prostatic hyperplasia (BPH) is a prevalent problem among older men, and its incidence is expected to increase as the human lifespan is prolonged. Symptoms of BPH, such as lower urinary tract symptoms (LUTS), have a negative impact on quality of life [1].

Prostate-specific antigen (PSA) is the most important prostate cancer screening tool and the majority of prostate cancers are detected with biopsy after abnormal PSA results. The ability to accurately detect prostate cancer can be compromised by any factor that decreases PSA. Several studies have found that obese men have lower PSA values than do nonobese men [2-7]. Because of hemodilution by the large plasma volume in obese men, some investigators have hypothesized that the PSA value is underestimated in obesity [8].

Multiple studies have reported that obese men have a larger prostate volume (PV) [9-13], and recently some studies have also revealed that a relationship exists between obesity and LUTS [9,11,14,15]. There have been few studies, however, concerning the effect of obesity on LUTS as

well as BPH parameters in Korean men, especially symptomatic BPH patients. Furthermore, we think it might be meaningful to investigate this correlation in a single institution. Therefore, we examined the correlation between body mass index (BMI) as a marker of obesity and PSA, PV, and International Prostate Symptom Score (IPSS) in symptomatic BPH patients.

MATERIALS AND METHODS

From January 2008 to December 2008, we retrospectively investigated 258 patients diagnosed with BPH in our institution. All patients underwent detailed clinical evaluations with the IPSS questionnaire and blood tests including PSA values. Transrectal ultrasound (TRUS) of the prostate was also performed. Anthropometric measurements including height and weight were performed. BMI was calculated as weight in kilograms divided by the square of height in meters (kg/m²). IPSS was used as the objective scale for degree of LUTS.

BPH was defined as a PV \geq 25 ml and IPSS \geq 8, which are commonly used as clinical BPH criteria [16]. BMI (kg/m²) groups were categorized according to the WHO BMI criteria for Asians as follows: underweight (BMI < 18.5), normal weight (BMI 18.5-23.0), overweight (BMI 23.0-27.5), and obese (BMI \geq 27.5) [17].

The exclusion criteria of this study were the use of medications affecting prostate growth, such as $5 - \alpha$ -reductase inhibitors and antiandrogens; the presence of neurogenic bladder dysfunction; confirmed prostate cancer by needle biopsy of the prostate; acute or chronic urinary retention status; a history of recurrent urinary tract infection or bladder stones; acute or chronic prostatitis within the previous 3 months; and a previous surgical procedure related to BPH.

Statistical analyses were performed by using the SPSS ver. 17.0 (SPSS Inc., Chicago, IL, USA) PSA values were natural-log-transformed to improve normality and to control for age when presenting the results because of the known association between PSA and age [18]. Thus, mean values of PSA are presented as age-adjusted (geometric) means in the figure but are referenced as mean PSA in the text, including tables, to simplify matters. One-way analysis of variance (ANOVA) was used to determine the statistical significance of the differences in age, PSA, PV, and IPSS among the groups according to BMI. Univariate analysis by Pearson's correlation coefficient was used to test the linearity of the relationships between PSA, PV, IPSS, and BMI. A p-value of < 0.05 was considered statistically significant.

RESULTS

1. Subjects characteristics

The mean age of the patients was 65.19 ± 9.13 years and their mean BMI was 23.7 ± 4.4 kg/m². Of the subjects,11 (4.3%) were diagnosed with hypertension and 19 (7.4%) with diabetes mellitus. All of the patients had been using medication and had well-controlled blood pressure or plasma glucose levels. No subjects had diagnosed metabolic syndrome.

The BMI distribution of the 258 subjects was as follows: 40, 72, 85, and 61 in the underweight (15.5%), normal weight (27.9%), overweight (32.9%), and obese (23.7%) groups,

 TABLE 1. Mean BMI, PSA and IPSS including subscores according to age group (n=258)

Age (yr)	BMI (kg/m ²)	PSA (ng/ml)	PV (ml)	IPSS (Obstructive/Irritative)		
50-59 (n=61)	23.71 ± 5.36	2.45 ± 1.49	41.27±19.34	$18.32{\pm}6.07 \\ (10.14{\pm}2.19/5.51{\pm}2.04)$		
60-69 (n=114)	23.96 ± 4.12	2.87 ± 1.83	48.37±26.92	$\begin{array}{c} 19.59{\pm}5.76 \\ (12.51{\pm}2.63/7.02{\pm}3.29) \end{array}$		
70-79 (n=83)	23.24±3.92	3.27 ± 2.06	55.61 ± 23.42	$\begin{array}{c} 21.66{\pm}6.63 \\ (14.05{\pm}3.25/9.53{\pm}2.86) \end{array}$		

PSA: prostate-specific antigen, BMI: body mass index, PV: prostate volume, IPSS: International Prostate Symptom Score

TABLE 2. Comparison of	f the stu	ıdy subject	s by BMI
------------------------	-----------	-------------	----------

	Mean±SD				
	Underweight (n=40)	Normal weight (n=72)	Overweight (n=85)	Obese (n=61)	p-value
Age (yr)	62.10 ± 11.07	68.26 ± 9.35	66.35 ± 7.19	63.16 ± 8.82	0.753
PSA (ng/ml)	3.42 ± 1.53	3.07 ± 1.88	2.74 ± 1.75	2.60 ± 1.44	0.023^{a}
PV (ml)	35.40 ± 12.82	46.78 ± 24.34	49.53 ± 27.14	59.95 ± 26.27	0.007^{a}
IPSS	15.13 ± 3.96	17.90 ± 5.98	20.19 ± 6.77	25.07 ± 5.22	0.023^{a}
Subscore of obstructive symptom	9.87 ± 2.45	11.04 ± 3.88	12.74 ± 4.52	15.39 ± 3.76	0.043^{a}
Subscore of irritative symptom	5.26 ± 1.68	6.86 ± 1.96	7.45 ± 3.29	9.68 ± 2.84	0.039^{a}

BMI: body mass index, PSA: prostate-specific antigen, PV: prostate volume, IPSS: International Prostate Symptom Score, ^a: p < 0.05

respectively. Mean PSA, BMI, PV, and IPSS including subscores according to each age group are shown in Table 1. Shown in Table 2 are the comparisons of the study subjects in each BMI group.

2. Prostate-specific antigen

The mean PSA value was 3.42 ± 1.53 , 3.07 ± 1.88 , 2.74 ± 1.75 , and 2.60 ± 1.44 ng/ml in each BMI group. The PSA value was lowest in the obese group. There were significant differences in PSA values among the BMI groups (p=0.023) (Table 2).

3. Prostate volume

The mean PV was 35.40 ± 12.82 , 46.78 ± 24.34 , 49.53 ± 27.14 , and 59.95 ± 26.27 ml in each BMI group. PV was largest in the obese group. There were significant differences in PV among the BMI groups (p=0.007) (Table 2).

4. International Prostate Symptom Score

The mean IPSS was 15.13 ± 3.96 , 17.90 ± 5.98 , 20.19 ± 6.77 , and 25.07 ± 5.22 in each BMI group. The IPSS and subscores for obstructive and irritative symptoms were highest in the obese group. There were significant differences in IPSS among the BMI groups (p < 0.05) (Table 2).

5. Associations between PSA, PV, IPSS and BMI

We examined the relationships between PSA, PV, IPSS,



and BMI. In the univariate analysis by Pearson's correlation coefficient, BMI correlated negatively with PSA (p=0.023) and positively with PV (p=0.001) and IPSS (p=0.02). The correlation coefficients of PSA, PV, and IPSS were -0.142, 0.320, and 0.470, respectively (Fig. 1).

DISCUSSION

Over the past decades, many different groups have investigated the influence of obesity on the development of BPH with conflicting results [19-23]. Freedland et al examined the association between BMI and PSA among men who underwent radical prostatectomy for prostate cancer. They found no association between BMI and PSA [24]. However, Bañez et al examined the association between obesity-related plasma hemodilution and PSA concentration among men who underwent radical prostatectomy for prostate adenocarcinoma. They suggested that the PSA value was underestimated in obesity and that lower PSA values were largely due to hemodilution by the large plasma volume in obese men [8]. Sohn et al investigated the association between BMI and PSA among 26,912 Korean men who visited health promotion centers. They noted that BMI was inversely correlated with PSA [25]. In this study, the mean PSA value was lowest in the obese group. In the univariate analysis by Pearson's correlation coefficient, BMI corre-



FIG. 1. Univariate analysis by Pearson's correlation coefficient between BMI and PSA, PV, and IPSS. (A) Correlation between BMI and PSA (Pearson's correlation coefficient= -0.142, p= 0.023), (B) Correlation between BMI and PV (Pearson's correlation coefficient=0.320, p=0.001), (C) Correlation between BMI and IPSS (Pearson's correlation coefficient=0.470, p=0.02). BMI: body mass index, PSA: prostate-specific antigen, PV: prostate volume, IPSS: International Prostate Symptom Score.

lated negatively with PSA (p=0.023).

In terms of correlation between PV and BMI, Kim et al investigated the relationship of PV with metabolic and anthropometric parameters. They reported that PV correlated positively with weight and height, but there was no statistical correlation between PV and BMI in the multivariable linear regression analysis [26]. A recent study of 465 men recruited through a health promotion center showed that PV was positively correlated with central obesity, as represented by waist circumference, but not with overall obesity, as represented by BMI [27]. By contrast, a US study of men who had undergone radical prostatectomy reported that BMI was positively associated with PV in those younger than 63 years [24]. In another study of men shown by biopsy to be without prostate cancer, BMI was directly associated with PV [28].

Obesity may influence prostatic enlargement and may also worsen urinary obstructive symptoms by increasing activity of sympathetic nervous systems [29]. Kristal et al examined several modifiable lifestyle factors related to the development of symptomatic BPH in 5,600 men enrolled in the placebo arm of the Prostate Cancer Prevention Trial who were followed for 7 years [19]. They reported significant increases in symptomatic BPH (IPSS>14) with obesity. Therefore, they suggested that obesity in adulthood was associated with a higher prevalence of LUTS. Rohrmann et al investigated the association between obesity and LUTS in the National Health and Nutrition Examination Survey (NHANES) III cohort [14]. They recognized that an increase in BMI after age 25 was positively associated with LUTS.

We examined patients who visited the department of urology and were diagnosed with BPH. The results of our study showed that the mean value of PV and IPSS increased with elevated BMI. In the correlation analysis, BMI correlated positively with PV (p=0.001) and IPSS (p=0.02). The correlation coefficients for PV and IPSS were 0.320 and 0.470, respectively.

Several recent studies concerning the relationships between BMI and BPH parameters have been reported, but these studies included healthy populations of men who visited a health promotion center. In the present study, we studied symptomatic BPH patients who visited the department of urology for evaluation or treatment. It was an aim of our study to reveal the relationship between BMI and BPH parameters in symptomatic BPH patients. Therefore, we think that it might be meaningful to announce the results of this study to BPH patients.

CONCLUSIONS

Patients with a large BMI tended to have a lower PSA value, larger PV, and higher IPSS. We hypothesize that weight loss could help in the relief of LUTS in BPH patients. We also note that the PSA value was underestimated in obesity. To prevent delay in early diagnosis of coexisting prostate cancer in obese patients, a study of prostate cancer screening in coordination with BMI is warranted.

Conflicts of Interest

The authors have nothing to disclose.

REFERENCES

- Rohrmann S, Smit E, Giovannucci E, Platz EA. Associations of obesity with lower urinary tract symptoms and noncancer prostate surgery in the Third National Health and Nutrition Examination Survey. Am J Epidemiol 2004;159:390-7.
- Fowke JH, Signorello LB, Chang SS, Matthews CE, Buchowski MS, Cookson MS, et al. Effects of obesity and height on prostate-specific antigen (PSA) and percentage of free PSA levels among African-American and Caucasian men. Cancer 2006;107: 2361-7.
- 3. Rodriguez C, Patel AV, Calle EE, Jacobs EJ, Chao A, Thun MJ. Body mass index, height, and prostate cancer mortality in two large cohorts of adult men in the United States. Cancer Epidemiol Biomarkers Prev 2001;10:345-53.
- Vollmer RT, Humphrey PA. Tumor volume in prostate cancer and serum prostate-specific antigen: analysis from a kinetic viewpoint. Am J Clin Pathol 2003;119:80-9.
- Dubois D, Dubois EF. A formula to estimate the approximate surface area if height and weight be known. Arch Intern Med 1916; 17:863-71.
- 6. Boer P. Estimated lean body mass as an index for normalization of body fluid volumes in humans. Am J Physiol 1984;247:F632-6.
- Garza C, Haas J, Himes J, Pradilla A, Raman L, Seidell J, et al. Physical status: the use and interpretation of anthropometry: report of a WHO Expert Committee. World Health Organ Tech Rep Ser 1995;854:1-452.
- Bañez LL, Hamilton RJ, Partin AW, Vollmer RT, Sun L, Rodriguez C, et al. Obesity-related plasma hemodilution and PSA concentration among men with prostate cancer. JAMA 2007;298:2275-80.
- Hammarsten J, Högstedt B, Holthuis N, Mellström D. Components of the metabolic syndrome-risk factors for the development of benign prostatic hyperplasia. Prostate Cancer Prostatic Dis 1998;1:157-62.
- Gupta A, Gupta S, Pavuk M, Roehrborn CG. Anthropometric and metabolic factors and risk of benign prostatic hyperplasia: a prospective cohort study of Air Force veterans. Urology 2006;68: 1198-205.
- Ozden C, Ozdal OL, Urgancioglu G, Koyuncu H, Gokkaya S, Memis A. The correlation between metabolic syndrome and prostatic growth in patients with benign prostatic hyperplasia. Eur Urol 2007;51:199-203.
- 12. Xie LP, Bai Y, Zhang XZ, Zheng XY, Yao KS, Xu L, et al. Obesity and benign prostatic enlargement: a large observational study in China. Urology 2007;69:680-4.
- Putnam SD, Cerhan JR, Parker AS, Bianchi GD, Wallace RB, Cantor KP, et al. Lifestyle and anthropometric risk factors for prostate cancer in a cohort of Iowa men. Ann Epidemiol 2000;10: 361-9.
- 14. Rohrmann S, Smit E, Giovannucci E, Platz EA. Association between markers of the metabolic syndrome and lower urinary tract symptoms in the Third National Health and Nutrition Examination Survey (NHANES III). Int J Obes (Lond) 2005;29:310-6.
- 15. Joseph MA, Harlow SD, Wei JT, Sarma AV, Dunn RL, Taylor JM, et al. Risk factors for lower urinary tract symptoms in a pop-

ulation-based sample of African-American men. Am J Epidemiol 2003;157:906-14.

- Hald T. Urodynamics in benign prostatic hyperplasia: a survey. Prostate Suppl 1989;2:69-77.
- 17. WHO Expert Consultation. Appropriate body-mass index for Asian populations and its implications for policy and intervention strategies. Lancet 2004;363:157-63.
- Werny DM, Thompson T, Saraiya M, Freedman D, Kottiri BJ, German RR, et al. Obesity is negatively associated with prostate-specific antigen in U.S. men, 2001-2004. Cancer Epidemiol Biomarkers Prev 2007;16:70-6.
- Kristal AR, Arnold KB, Schenk JM, Neuhouser ML, Weiss N, Goodman P, et al. Race/ethnicity, obesity, health related behaviors and the risk of symptomatic benign prostatic hyperplasia: results from the prostate cancer prevention trial. J Urol 2007;177: 1395-400.
- Laven BA, Orsini N, Andersson SO, Johansson JE, Gerber GS, Wolk A. Birth weight, abdominal obesity and the risk of lower urinary tract symptoms in a population based study of Swedish men. J Urol 2008;179:1891-5.
- Park HK, Lee HW, Lee KS, Byun SS, Jeong SJ, Hong SK, et al. Relationship between lower urinary tract symptoms and metabolic syndrome in a community-based elderly population. Urology 2008;72:556-60.
- 22. Koo KC, Cho KS, Kang EM, Kwon SW, Hong SJ. The relationship between metabolic syndrome and prostate volume in men over

sixties who underwent prostate health check-up. Korean J Urol 2008;49:813-7.

- 23. Lee SH, Kim JC, Lee JY, Kim JH, Oh CY, Lee SW, et al. Effects of obesity on lower urinary tract symptoms in Korean BPH patients. Asian J Androl 2009;11:663-8.
- Freedland SJ, Platz EA, Presti JC Jr, Aronson WJ, Amling CL, Kane CJ, et al. Obesity, serum prostate specific antigen and prostate size: implications for prostate cancer detection. J Urol 2006; 175:500-4.
- 25. Sohn JC, Lim MS, Chang HS, Park CH, Kim CI. The association of body mass index and prostate-specific antigen. Korean J Urol 2007;48:1121-4.
- Kim YD, Yang WJ, Song YS, Park YH. Correlation between prostate volume and metabolic or anthropometric factors in male visitors to a health promotion center. Korean J Urol 2008;49:139-44.
- Kim GW, Doo SW, Yang WJ, Song YS. Effects of obesity on prostate volume and lower urinary tract symptoms in Korean men. Korean J Urol 2010;51:344-7.
- Ochiai A, Fritsche HA, Babaian RJ. Influence of anthropometric measurements, age, and prostate volume on prostate-specific antigen levels in men with a low risk of prostate cancer. Urology 2005;66:819-23.
- Dahle SE, Chokkalingam AP, Gao YT, Deng J, Stanczyk FZ, Hsing AW. Body size and serum levels of insulin and leptin in relation to the risk of benign prostatic hyperplasia. J Urol 2002;168: 599-604.