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Changes in health-related quality of life after radical prostatectomy for prostate cancer: A longitudinal cohort study in Korea

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Purpose: To evaluate longitudinal changes in health-related quality of life and satisfaction after radical prostatectomy for the treatment of prostate cancer in Korean men.

Materials and Methods: Data from patients with localized prostate cancer who underwent radical prostatectomy between January 2012 and December 2013 were analyzed. Patients completed the Expanded Prostate Cancer Index Composite questionnaire at scheduled clinic visits (baseline and 1, 3, 6, and 12 months post-surgery).

Results: Data from 211 men were evaluated. Urinary domain summary scores decreased postoperatively but improved over the 12-month follow-up period. Urinary function and incontinence had not recovered to baseline values at 12 months, whereas urinary bother and irritation/obstruction scores showed no differences from baseline at 12 months. Sexual function improved at each timepoint to 12 months but did not recover to baseline values; sexual bother was approximately 50% of baseline values at 12 months. Recovery of sexual function was significantly greater in patients who underwent robotic surgery versus those who underwent open surgery. At the end of the study period, 66.3% of patients were satisfied with the treatment they had received; multivariate analysis showed that only urinary function recovery was significantly associated with overall satisfaction.

Conclusions: Urinary outcomes and sexual outcomes had not returned to baseline at 12 months. Recovery of sexual function was significantly greater in patients who underwent robotic surgery. Only recovery of urinary function was significantly associated with overall satisfaction 12 months after radical prostatectomy in this cohort of Korean men.

Keywords: Prostatectomy; Prostatic neoplasms; Quality of life; Surveys and questionnaires

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INTRODUCTION

Radical prostatectomy is a key approach to the treatment of localized prostate cancer. The procedure is associated with favorable survival, highlighting the importance of health-related quality of life (HRQOL) and functional outcomes after surgery [1-3]. Urinary incontinence and erectile dysfunction are common adverse events and may have a profound impact on HRQOL in patients with prostate cancer. Patients with clinically localized prostate cancer have multiple treatment options and the risk:benefit balance is an important aspect in treatment decision-making

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[4]. Two recent population-based prospective cohort studies have evaluated HRQOL after contemporary treatments for localized prostate cancer, showing a distinct pattern of adverse effects associated with each treatment strategy [5,6].

The incidence of prostate cancer has increased steadily over the last decade in East Asia, most significantly in Korea [7]. Although a number of studies have examined HRQOL after radical prostatectomy in Western and Japanese populations [8], to date, none have evaluated Korean prostate cancer patients. The heterogeneity of prostate cancer has been documented in patients from different geographical and ethnic populations [9]. Race and ethnicity are also important factors in HRQOL assessment as they are known to impact on satisfaction with care [10].

In this study, we have prospectively evaluated longitudinal changes in HRQOL in Korean prostate cancer patients after radical prostatectomy. In addition, we have attempted to identify factors that may impact on overall satisfaction after treatment.

MATERIALS AND METHODS

1. Study population

Data from patients who underwent radical prostatectomy at Asan Medical Center between January 2012 and December 2013 were reviewed. All surgery was conducted by a single, highly experienced surgeon. Patients were required to complete the Expanded Prostate Cancer Index Composite (EPIC) questionnaire at baseline and post-surgery. Patients who received neoadjuvant or adjuvant therapy and those with inadequate clinical data or incomplete questionnaires were excluded. The study protocol was approved by the Institutional Review Board of Asan Medical Center (approval number: 2012-0036) and informed consent was obtained from each patient.

2. Measurement of HROOL

Patients were routinely scheduled to visit the clinic at baseline and at 1, 3, 6, and 12 months post-surgery. At each visit, patients were asked to complete the Korean version of the EPIC [11] questionnaire on paper in the waiting room before meeting the urologist.

The EPIC questionnaire consists of 50 questions and is divided into four domains: bowel, urinary, sexual, and hormonal. The four domain summary scores and subscale scores were calculated on a scale of 0–100, with higher scores representing better HRQOL [12].

Overall satisfaction was determined based on a single question 'Overall, how satisfied are you with the treatment

you received for your prostate cancer?' Patients who responded that they were 'extremely satisfied' or 'satisfied' were classified as the 'satisfied' group.

3. Statistical analysis

Paired t-tests were used to assess differences in subscale scores at each timepoint and at different timepoints along the same domain or subscale. Repeated-measures ANOVA was used to compare HRQOL changes over time between patients undergoing robot-assisted laparoscopic prostatectomy (RALP) and those undergoing retropublic radical prostatectomy (RRP). Multivariate analysis with a stepwise backwards elimination approach was performed to identify significant influencing factors for overall satisfaction after treatment. All statistical tests were two-tailed with a significance level of 0.05; analyses were

Table 1. Patient characteristics (n=211)

Characteristic	Value
Age (y)	64.7
Residential area	
Metropolitan	61 (28.9)
Urban	84 (39.8)
Suburban/rural	66 (31.3)
Educational level	
≥College	85 (40.3)
High school	69 (32.7)
≤Middle school	57 (27.0)
Charlson comorbidity index	
0	133 (63.0)
1	44 (20.9)
2	26 (12.3)
≥3	8 (3.8)
Mean prostate-specific antigen (ng/mL)	10.5
Mean prostate volume (mL)	35.0
Clinical T stage	
T1	114 (54.0)
T2	88 (41.7)
T3	9 (4.3)
Pathologic T stage	
T2	136 (64.4)
T3a	55 (26.1)
T3b	20 (9.5)
Surgical approach	
Open	43 (20.4)
Robot-assisted	168 (79.6)
Pelvic node dissection	105 (49.8)
Nerve sparing	
Unilateral	53 (25.1)
Bilateral	158 (74.9)

Values are presented as mean only or number (%).



performed using IBM SPSS Statistics ver. 21.0 software (IBM Co., Armonk, NY, USA).

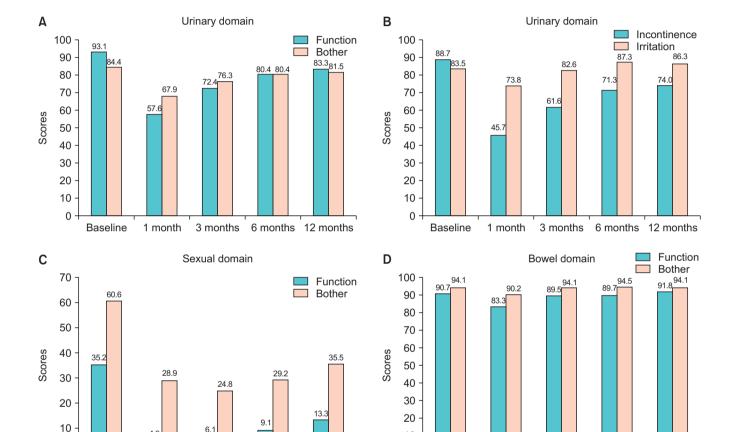
RESULTS

Data from a total of 211 patients were included in the analyses. Each patient was evaluated at five visits; therefore, a total of 1,055 EPIC questionnaires and medical records were analyzed. Patient characteristics are summarized in Table 1; the mean patient age was 64.7 years, mean prostate-

specific antigen level was 10.5 ng/mL, and mean prostate volume was 35.0 mL. Of the 211 patients, 136 (64.4%) had organ-confined prostate cancer and 168 (79.6%) underwent robot-assisted radical prostatectomy.

1. Urinary domain

Urinary domain summary scores are shown in Fig. 1A. Prior to surgery, mean urinary function scores were higher than urinary bother scores. However, at 1 month, urinary function had decreased to a greater extent than



10

0

Baseline

Ε Hormone domain Function Bother 100 87.4_90.8 89.0<u>91.</u>3 87.9 88 90 81.6 80 70 60 50 40 30 20 10 6 months 12 months Baseline 1 month 3 months

3 months

6 months

12 months

Fig. 1. Longitudinal health-related quality of life domain scores. (A) Urinary function and bother score. (B) Urinary incontinence and irritation/obstruction score. (C) Sexual function and bother score. (D) Bowel function and bother score. (E) Hormone function and bother score.

1 month

3 months

0

Baseline

1 month

6 months 12 months



urinary bother. Urinary function improved at 3 months and reached a new baseline at 6 months. Urinary function had not recovered fully at 12 months, whereas urinary bother showed no difference from baseline at 12 months.

In the EPIC questionnaire, the urinary domain is separated into two distinct subscales, incontinence and irritation/obstructive (Fig. 1B). At 1 month post-surgery, mean urinary incontinence had decreased by a greater degree than mean irritation/obstruction. Although mean urinary incontinence had improved at 12 months, values did not return to baseline. By contrast, following a slight decrease at 1 month, mean urinary irritation/obstruction returned to baseline at 3 months and then rose above baseline from 6 months onwards. Medical records showed that 14.2% of patients were incontinent at 12 months post-surgery in this study cohort.

2. Sexual domain

Post-surgery sexual domain scores showed the greatest reduction of all four domains. At baseline, mean sexual function was lower than mean sexual bother (Fig. 1C). At 1 month, the mean sexual function score had decreased greatly from baseline. Gradual improvement was seen at each time point but scores had not recovered by 12 months. The mean sexual bother score had decreased by >50% at 1 month post-surgery and remained low at 12 months.

3. Bowel domain

Mean bowel function decreased slightly at 1 month postsurgery and returned to baseline levels at 3 months (Fig. 1D). Mean bowel function and mean bowel bother showed no significant differences from baseline at any postoperative timepoint.

4. Hormone domain

After a slight decrease at 1 month, mean hormone function increased above baseline at 3 months (Fig. 1E). Mean hormone bother also decreased at 1 month but returned to baseline at 3 months post-surgery. No statistically significant differences in mean hormone function and mean hormone were seen at any postoperative timepoint.

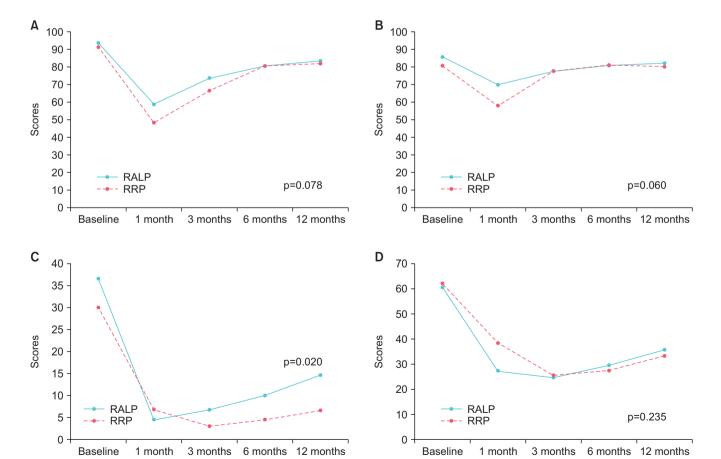


Fig. 2. Comparison of urinary and sexual outcomes between robot-assisted and open radical prostatectomy groups. (A) Urinary function. (B) Urinary bother. (C) Sexual function. (D) Sexual bother. Based on repeated-measures ANOVA. Statistical tests for interaction of surgery group and time. RALP, robot-assisted laparoscopic prostatectomy; RRP, retropubic radical prostatectomy.



5. Comparison of urinary and sexual outcomes in patients undergoing robot-assisted or open radical prostatectomy

At baseline, there were no significant differences in age, urinary function, urinary bother, sexual function, and sexual bother between the RALP and RRP groups. Repeated-measures ANOVA showed no significant changes over time in terms of urinary function (p=0.078; Fig. 2A), urinary bother (p=0.060; Fig. 2B), or sexual bother (p=0.235; Fig. 2D). However, changes in sexual function over time were significantly different between the two groups (p=0.020; Fig. 2C) and recovery of sexual function was significantly better in the RALP group compared with the RRP group from 3-12 months post-surgery.

6. Overall satisfaction

At 1, 3, 6, and 12 months after radical prostatectomy, 68.7%, 65.0%, 64.5%, and 66.3% of patients indicated that they were satisfied with the treatment they had received (Table 2). Multivariate analysis showed that only recovery

Table 2. Overall satisfaction 12 months after radical prostatectomy

Variable —	No. of patients satisfied with treatment			
	1 month	3 months	6 months	12 months
Overall	145 (68.7)	137 (65.0)	136 (64.5)	140 (66.3)
RALP	120 (71.4)	109 (64.9)	106 (63.1)	112 (66.7)
RRP	25 (58.1)	28 (65.1)	30 (69.7)	28 (65.1)

Values are presented as number (%).

RALP, robot-assisted laparoscopic prostatectomy; RRP, retropubic radical prostatectomy.

Table 3. Univariable and multivariable logistic regression analyses to identify which EPIC domain subscale changes are associated with overall satisfaction 12 months post-surgery

Variable	Univariate		Multivariat	Multivariate	
variable	OR (95% CI)	p-value	OR (95% CI)	p-value	
Age	1.014 (0.965–1.065)	0.587			
Residential area					
Metropolitan	Reference				
Urban	0.854 (0.382–1.907) 0.699				
Suburban/rural	0.520 (0.228–1.188) 0.121				
Educational level					
≥College	Reference				
High school	0.797 (0.380-1.670)	0.547			
≤Middle school	0.544 (0.252-1.174)	0.121			
Charlson comorbidity index					
0	Reference				
1	0.871 (0.409-1.854)	0.720			
2	0.488 (0.148-1.608)	0.238			
≥3	1.220 (0.227-6.562)	0.817			
RALP (vs. RRP)	1.083 (0.440-2.667)	0.863			
EPIC domain subscale ^a					
Urinary function	1.037 (1.015–1.059)	0.001	1.038 (1.006-1.071)	0.021	
Urinary bother	1.026 (1.007-1.046)	0.009	0.992 (0.962-1.024)	0.633	
Sexual function	1.010 (0.994-1.027)	0.203			
Sexual bother	1.005 (0.995-1.015)	0.342			
Bowel function	1.021 (0.994–1.050)	0.132			
Bowel bother	1.046 (1.010-1.084)	0.012	1.038 (0.999-1.079)	0.059	
Hormone function	1.013 (0.991-1.035)	0.239			
Hormone bother	1.019 (0.997-1.041)	0.095			

EPIC, Expanded Prostate Cancer Index Composite; OR, odds ratio; CI, confidence interval; RALP, robot-assisted laparoscopic prostatectomy; RRP, retropubic radical prostatectomy.

^a:Each EPIC domain subscale variable represents a change from baseline to 12 months post-surgery.



of urinary function was significantly associated with overall satisfaction 12 months after radical prostatectomy (Table 3).

DISCUSSION

In this study, Korean patients with prostate cancer reported a more pronounced decline in sexual function and bother after radical prostatectomy, compared with other HRQOL outcomes. However, sexual outcome was not associated with overall satisfaction and an association was seen only with the recovery of urinary function.

Overall, the pattern of change related to each HRQOL domain in this cohort of Korean men was similar to that reported in previous studies [2,13-17]. However, some differences were apparent. In comparison with a study of American men with prostate cancer [14], the EPIC sexual function score was approximately twice as high as in our study, recovering to half of the baseline values at 12 months post-surgery. However, in our study, the sexual function score was lower at baseline and recovered to approximately one third of baseline values at 12 months post-surgery, which is similar to Japanese study results [16,17].

The most likely cause of the observed differences in sexual outcomes versus Western studies is ethnicity. Namiki et al. [10] investigated sexual function following radical prostatectomy in a prospective longitudinal study of Japanese and American men. In the Namiki et al.'s study [10], Japanese men reported lower sexual function scores at baseline and American men were more likely than Japanese men to regain their baseline sexual function. In this study, the mean baseline sexual function score was lower than that of Western studies [13,14]. In a primary care study of Korean men [18], the prevalence of erectile dysfunction was 325%, 52.4%, and 68.5% in men aged 40-49, 50-59, and ≥60 years, respectively, which is higher than that seen in other countries, including Japan [10,19]. In a sub-analysis of preoperatively potent patients in this study cohort, sexual function and bother increased from 6 months after surgery and reached approximately 80% of baseline at 12 months, a result similar to that seen in the American study (Supplementary Fig). In these preoperatively potent patients, the potency rate at 12 months post-surgery was 53.5%. Moreira et al. [20] reported that only 2% of men in Korea had consulted a doctor about their sexual problems. This is largely due to the belief that the problem is not sufficiently serious and therefore not of significant concern. In Korean men, such perception of sexual problems may have affected their sexual function and recovery after surgery.

HRQOL can be dependent on treatment modality.

Because of the difficulties associated with conducting randomized trials, comparing outcomes related to RALP and RRP is challenging and studies have shown mixed results [21-24]. A Cochrane review [25], which is the most recent study on this issue, found no evidence supporting the effectiveness of RALP (in terms of oncological and functional outcome) over RRP. However, men undergoing RALP had shorter hospital stays and lower rates of blood transfusion. Consistent with this, we observed that the mean hospital stay was shorter (8.3 days vs. 11.9 days, p<0.001) and blood transfusion rates lower (2.4% vs. 9.3%, p<0.001) after RALP than after RRP. Although the uptake of RALP in East Asia has been somewhat slower and less widespread than in Western countries, the use of RALP has been steadily increasing over the past decade in East Asia, including Korea [7,26]. However, to our knowledge, there have been no published studies comparing postoperative longitudinal HRQOL changes in Asian men (including Korean patients) with prostate cancer undergoing RALP and RRP. Our results showed that recovery of sexual function was greater in the RALP group than the RRP group, an observation that is consistent with some previous reports [23,24,27].

Previous studies have shown that 16% to 19% of patients with localized prostate cancer regret their treatment choice [28,29]. In these studies, sociodemographic factors and HRQOL outcomes impacted on satisfaction and regret after treatment. Notably, the overall satisfaction rate 12 months after radical prostatectomy in Korean men was 66.2%, which is considerably lower than that reported in the American population [28-30]. In the current study, we found that recovery of urinary function was the only influencing factor for overall satisfaction and the notable decline in sexual domain score, sexual function, and bother did not significantly affect overall satisfaction. As described above, perception of sexual problems may have contributed to these results.

We acknowledge several limitations in this study. First, the duration of follow-up was not sufficient to evaluate long-term results and functional outcomes have been reported to change after 2 or more years in several studies [3,14-16]. In our study cohort, we found that the potency rates of preoperatively potent patients continuously increased from 53.5% at 1 year to 61.8% at 4 years after surgery. Secondly, treatment selection was not randomized and the study population may not be representative of all Korean men undergoing radical prostatectomy. Thirdly, although there were no preoperative differences in age and HRQOL domain scores in the RALP and RRP groups, baseline characteristics

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could not be fully adjusted. Fourthly, sociodemographic variables were not considered in the analysis of overall satisfaction after surgery. However, we believe the present study to be of value as a prospectively designed, single-surgeon, single-institution series of consecutive cases. Despite the short follow-up period, these data can be considered to be meaningful because changes in HRQOL are most marked during the first year post-surgery. The findings are also valuable in terms of the consistency and quality of the data because the study included only those patients who completed the questionnaire at all timepoints. In addition, to our knowledge, this is the first study to prospectively compare patient-reported HRQOL changes between RALP and RRP in Asian men. Nonetheless, additional multicenter, randomized studies are required to confirm our observations.

CONCLUSIONS

In this cohort of Korean men undergoing radical prostatectomy, urinary function and incontinence had not recovered to baseline values at 12 months, whereas urinary bother and irritation/obstruction recovered fully by the end of the study period. Sexual outcomes declined significantly after surgery and did not fully recover.

Recovery of sexual function was significantly better in the patient group who underwent robotic surgery compared with those who underwent open surgery. Only urinary function recovery was significantly associated with overall satisfaction 12 months after radical prostatectomy.

CONFLICTS OF INTEREST

The authors have nothing to disclose.

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SUPPLEMENTARY MATERIAL

Scan this QR code to see the supplementary material, or visit https://www.icurology.org/src/sm/icurology-59-313-s001.pdf.



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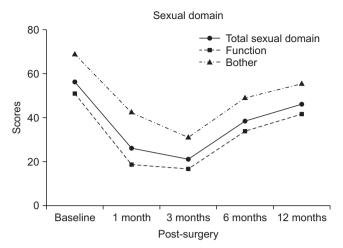
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Supplementary Fig. Longitudinal sexual domain scores in preoperatively potent patients.