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Appropriate preoperative membranous urethral length predicts recovery of urinary continence after robot-assisted laparoscopic prostatectomy

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Abstract

Purpose: We investigated that preoperative membranous urethral length (MUL) would be associated with the recovery of urinary continence after robot-assisted laparoscopic prostatectomy (RALP).

Patients and methods: We studied 204 patients who underwent RALP between May 2013 and March 2016. All patients underwent pelvic magnetic resonance imaging (MRI) preoperatively to measure MUL. Urinary continence was defined as the use of one pad or less (safety pad). The 204 patients were divided into two groups: continence group, those who achieved recovery of continence at 3, 6, and 12 months after RALP, and incontinence group, those who did not. We retrospectively analyzed the patients in terms of preoperative clinical factors including age, body mass index (BMI), estimated prostate volume, neurovascular bundle salvage, history of preoperative hormonal therapy, and MUL.

Results: The safety pad use rate was 69.6%, 86.9%, and 91.1% at 3, 6, and 12 months, respectively. On univariate and multivariate analyses, MUL were significant factors in every term of recovery of urinary continence in both groups. According to the receiver operating characteristic (ROC) curve analysis, the preoperative MUL that could best predict early recovery of urinary continence at 3 months after RALP was 12 mm.

Conclusions: We suggest that preoperative MUL > 12 mm would be a predictor of early recovery of urinary continence after RALP.

Keywords: Membranous urethral length, Urinary continence, Robot-assisted laparoscopic prostatectomy

Introduction

Urinary incontinence is one of the most unfavorable complications influencing the quality of life for patients after radical prostatectomy (RP). In early 2000, the initial robot-assisted laparoscopic prostatectomy (RALP) was performed with the da Vinci surgical system [1]. Currently, RALP has become a more popular surgical procedure for RP in Japan. RALP is expected to achieve better outcomes regarding recovery of urinary continence than did the conventional procedure. Its advanced technology provides a three-dimensional operative view and laparoscopic instruments that mimic the movement

of the human wrist. For the robotic approach, a metaanalysis of 51 studies showed statistically significant improvement in urinary continence recovery at 12 months with RALP compared to retropubic and laparoscopic RP [2].

Predictive factors for recovery of urinary continence after RP, such as patient age, body mass index (BMI), and prostate volume, have been reported [2, 3]. Membranous urethral length (MUL) as measured by magnetic resonance imaging (MRI) also was reported to be a strong predictive factor for recovery of urinary continence in a systematic review and meta-analysis [4]. These reports demonstrated that preoperative MUL is associated significantly and positively with a return to continence following RP. However, few reports exist on the

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association of MUL and recovery of urinary continence after RALP in the Japanese population.

We evaluated the association of preoperative MUL with the recovery of urinary incontinence after RALP in Japanese patients.

Patients and methods

We performed RALP using the da Vinci Si surgical system in 204 consecutive patients from whom we could collect the Expanded Prostate Cancer Index Composite (EPIC) questionnaire [5] at least preoperatively and 3 months after RALP between May 2013 and March 2016. All patients underwent pelvic MRI preoperatively. Most patients had no urinary incontinence before surgery; only ten patients had incontinence more than once a week. All surgical procedures were performed by four surgeons (YK, MK, RT, and WO). All surgeons are over 10 years as urologists, and each surgeon experienced RALP in more than 40 cases. RALP was performed via the conventional transperitoneal approach using the four-armed da Vinci surgical robot system [6]. In all cases, we performed the Rocco technique for posterior reconstruction of Denonvillier's facia [7], anterior preservation [8], and bladder neck preservation [9] for preventing incontinence. Hemi-nerve sparing was performed depending on the cancer status [10]. Pelvic lymph node dissection also was performed in patients with a high risk of cancer according to the D'Amico criteria. We also offered all patients pelvic floor exercise education during the operative period.

The MUL was measured by T2-weighted coronal and sagittal sections as a distance from the prostatic apex to the level of the urethra at the penile bulb on preoperative pelvic MRI (Fig 1) [11]. In terms of measuring the MUL, a number of urologists evaluated MUL of each

case at the preoperative conference. Thereafter, the data of MUL was remeasured by a researcher.

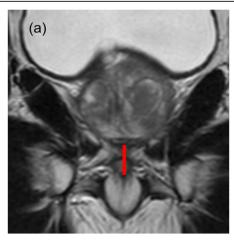
Urinary continence was evaluated at 3, 6, and 12 months after RALP using question 5 of the EPIC questionnaire [5]. Continence was defined as the use of one pad or less as a safety pad. We also examined the quality of life (QOL) score regarding urinary continence preand postoperatively using question 12 of the EPIC questionnaire [5].

The patients were divided into two groups: continence group, those who achieved recovery of urinary continence within 3, 6, 12 months, and incontinence group, those who did not.

Univariate analysis was performed with the t test, analysis of variance, chi-square, and Fisher's exact test between continence and incontinence group regarding preoperative clinical factors including patient age, BMI, estimated prostate volume, clinical stage, neurovascular bundle salvage, history of preoperative hormonal therapy, positive surgical margin, leakage at vesicourethral anastomosis, and MUL. Multivariate analysis was performed using a logistic regression model, and significances were tested using a likelihood ratio test. Statistical analyses were performed using JMP software (SAS Institute, Inc., Cary, NC, USA). For all statistical comparisons, differences with P < 0.05 were considered statistically significant.

Result

Mean patient age was 65 years, BMI was 23.7 kg/m², prostate-specific antigen (PSA) was 6.5 ng/ml, and MUL was 13.1 mm. Preoperative hormonal therapy was given in 34 (16.7%) patients, and hemi–nerve sparing was performed in 70 (34.6%; Table 1). The safety pad rate was 69.6%, 86.9%, and 91.1% at 3, 6, and 12 months, respectively, and the pad-free rate was 33.8%, 49.7%, and 64.3%, respectively. The QOL regarding the urinary condition



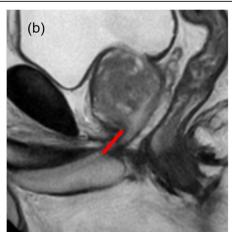


Fig. 1 The preoperative membranous urethral length as the distance from the prostatic apex to the level of the urethra at the penile bulb is measured in the T2-weighted MRI **a** coronal and **b** sagittal planes

Table 1 Patient characteristics

| Characteristic | Total ($n = 204$) |
|---|---------------------|
| Median (range) | |
| Observation period, days | 350 (82–545) |
| Age, years | 65 (41–76) |
| BMI, kg/m ² | 23.7 (17.2–37.6) |
| PSA level, ng/ml | 6.5(3.5-46.4) |
| Preoperative MUL, mm | 13.1 (4.5–22.9) |
| Estimated prostate volume, g | 38 (7–94) |
| Console time, min | 143 (87–351) |
| Operation time without console, min | 52 (18–93) |
| Intraoperative bleeding, ml* | 70 (10–1243) |
| N (%) | |
| Clinical stage | |
| ^T2a | 159 (77.9%) |
| T2b | 24 (11.8%) |
| ^T2c | 21 (10.3%) |
| Preoperative hormonal therapy history | 34 (16.7%) |
| Lymph node dissection | 58 (28.4%) |
| Neurovascular bundle saving | 70 (34.6%) |
| Positive surgical margin | 59 (28.9%) |
| Leakage at the vesicourethral anastomosis | 17 (8.3%) |

^{*}Including urine

after RALP was worst at 3 months. It improved gradually, and at 1 year postoperatively, it had improved to the preoperative status (Fig. 2).

In the univariate analysis for recovery of urinary continence 3 months after RALP, patient age (P = 0.034), and MUL (P < 0.001) were statistically significantly associated

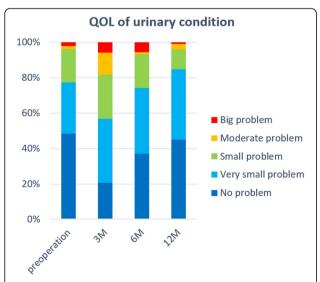


Fig. 2 QOL score of urinary condition per time period after RALP. The QOL status after RALP had the worst at 3 months. It was gradually improved, and 1 year after surgery was almost improved

with safety pad use. At 6 months after RALP, MUL (P = 0.004), console time, and leakage at vesicourethral anastomosis were statistically significant. At 12 months after RALP, MUL (P = 0.023) was statistically significant (Table 2). On multivariate analysis, patient age, leakage at vesicourethral anastomosis, and MUL achieved statistical significance with safety pad use. Among them, MUL was the most statistically significant with safety pad at 3 months after RALP (Table 3). Moreover, we estimated the optimal length of the MUL that could best classify between the continence and incontinence group. The cutoff point of MUL for recovery of urinary continence at 3 months after RALP was determined using the receiver operating characteristic (ROC) analysis. We identified a reasonable cutoff point of MUL to be 12 mm (Fig. 3). MUL > 12 mm was a favorable predictor of recovery of urinary continence at 3 months after RALP. Furthermore, using the cutoff point of MUL 12 mm, our cases were classified into continence and incontinence groups with a sensitivity of 80% and specificity of 70% (Fig. 3).

Discussion

RALP is expected to affect not only cancer control but also functional outcomes, such as continence and potency [12]. Especially, many reviews reported that patients who underwent RALP would tend to recover urinary continence within 1 year postoperatively earlier than patients who underwent retropubic RP (RRP) [12–14]. Early recovery of urinary continence is one of the strongest points of RALP. However, some patients suffer severe urinary incontinence after RALP. Therefore, we conducted this study to investigate factors that influence recovery of urinary continence in patients undergoing RALP.

Several preoperative predictive factors for recovery of urinary continence after RALP, such as age, body mass index, prostate size, medical comorbidities, history of transurethral resection of the prostate (TURP), and history of preoperative urinary continence or lower urinary tract symptoms, were identified as studied previously. MUL based on preoperative imaging also is a predictive factor associated with urinary continence [15]. Coakley et al. [16] reported the first study using endorectal MRI to measure preoperative MUL and showed a correlation of MUL with urinary continence after RRP [13]. They demonstrated that a longer preoperative MUL was associated with a faster recovery of continence; at 1 year postoperatively, 120 of 134 patients (89%) with preoperative MUL > 12 mm were completely continent, compared to only 35 of 46 (76%) whose preoperative MUL was ≤ 12 mm. Despite the preoperative median length of MUL seems to be a minor difference among continence and incontinence groups in our study, in prior studies, an increase in MUL as low as 1 mm can increase the odds of return to continence up to 200%

Table 2 Univariate analysis

| Parameters | Continence at 3 | months ($n = 204$ | Continence at 6 months ($n = 175$) | | | Continence at 12 months ($n = 112$) | | | |
|---|------------------------|-------------------------|--------------------------------------|------------------------|-------------------------|---------------------------------------|------------------------|-------------------------|-------|
| | Continence $(n = 142)$ | Incontinence $(n = 62)$ | Р | Continence $(n = 152)$ | Incontinence $(n = 23)$ | Р | Continence $(n = 102)$ | Incontinence $(n = 10)$ | Р |
| Median (range) | | | | | | | | | |
| Age, years | 65 (41–74) | 67 (51–76) | 0.034 | 66 (58–76) | 64 (41–75) | 0.989 | 66 (50–76) | 64 (58–73) | 0.689 |
| BMI, kg/m ² | 23.6 (17.2–37.6) | 23.9 (18.9–33.1) | 0.213 | 23.8 (17.2–37.6) | 23.6 (18.9–33.1) | 0.507 | 23.7 (17.2–32.7) | 24.5 (18.9–33.1) | 0.391 |
| PSA level, ng/ml | 6.7 (3.5-46.4) | 5.6 (4.2–24.1) | 0.178 | 6.4 (3.5–46.4) | 6.6 (4.2–14.5) | 0.833 | 6.2 (3.8–32.1) | 6.8 (4.2–16.7) | 0.669 |
| Preoperative MUL, mm | 13.6 (8.3–22.9) | 11.1 (4.5–17.9) | < .001 | 13.5 (6.5–22.9) | 11.1 (4.5–17.3) | 0.004 | 13.4 (6.5–22.9) | 10.8 (4.5–17.3) | 0.023 |
| Estimated prostate volume, g | 38 (7–94) | 38 (17–92) | 0.481 | 38 (7–94) | 41 (20–92) | 0.119 | 39 (14–94) | 49 (30–92) | 0.758 |
| Console time, min | 140 (87–351) | 150 (99–339) | 0.062 | 139 (89–253) | 157 (99–280) | 0.028 | 135 (89–253) | 177 (99–280) | 0.059 |
| Operation time without console, min | 51 (18–93) | 54 (23–86) | 0.792 | 49 (18–93) | 54 (23–86) | 0.477 | 51 (24–93) | 56 (23–86) | 0.506 |
| Intraoperative bleeding, ml | 68 (10–1243) | 83 (16–535) | 0.759 | 66 (14–1243) | 85 (20–397) | 0.696 | 70 (14–1243) | 78 (26–320) | 0.967 |
| N (%) | | | | | | | | | |
| Clinical stage | | | 0.218 | | | 0.763 | | | 0.532 |
| ^T2a | 109 (76.8%) | 50 (80.6%) | | 119 (78.3%) | 18 (78.2%) | | 83 (81.4%) | 9 (90%) | |
| T2b | 20 (14.1%) | 4 (6.5%) | | 19 (12.5%) | 2 (8.7%) | | 13 (12.7%) | 1 (10%) | |
| ^T2c | 13 (9.1%) | 8 (12.9%) | | 14 (9.2%) | 3 (13.1%) | | 6 (5.9%) | 0 | |
| Preoperative hormonal therapy history | 21 (14.8%) | 13 (20.9%) | 0.276 | 20 (13.2%) | 5 (21.7%) | 0.298 | 11 (10.8%) | 3 (30%) | 0.122 |
| Lymph node dissection | 40 (28.2%) | 18 (29%) | 0.901 | 43 (28.3%) | 6 (26.1%) | 0.825 | 25 (24.5%) | 2 (20%) | 0.746 |
| Neurovascular bundle saving | 52 (36.6%) | 18 (29%) | 0.312 | 57 (37.5%) | 8 (34.8%) | 0.901 | 39 (38.2%) | 4 (40%) | 0.951 |
| Positive surgical margin | 43 (30.3%) | 17 (27.4%) | 0.678 | 45 (29.6%) | 4 (17.4%) | 0.224 | 29 (28.4%) | 1 (10%) | 0.209 |
| Leakage at vesicourethral anastomosis | 9 (6.3%) | 9 (14.5%) | 0.068 | 8 (5.3%) | 7 (30.4%) | 0.007 | 5 (4.9%) | 2 (20%) | 0.059 |

BMI body mass index, PSA prostate-specific antigen, MUL membranous urethral length

[17]. Our study demonstrated that preoperative MUL > 12 mm was the strongest factor indicating recovery of urinary continence 3 months after RALP. We identified 3 months after RALP as the early period for recovery of urinary continence because the result of QOL testing at 3 months after RALP was by no means satisfactory, and there still was room for improvement.

A previous study reported that MUL has anatomical variation [14]. Our study showed similar data as those

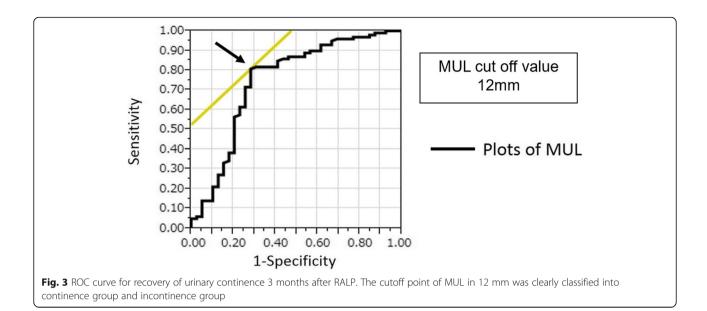
reported in Korean patients [18, 19]. The average MUL in Asian patients may be approximately 12 mm. On the other hand, MUL in reports from the USA and Europe is slightly longer than that of Asians, but the racial difference is not clear because the number of reports is too small [15, 20].

MUL also has been associated with urinary continence recovery after RP [4, 15, 17–19]. Paparerl et al. [11] reported that a loss ratio of MUL between pre- and postoperatively

Table 3 Multivariate analysis

| Parameters | Continence at 3 months ($n = 204$) | | | Continence at 6 months (n= | | 175) | Continence at 12 months ($n = 112$) | | | |
|---------------------------------------|--------------------------------------|-----------|---------|----------------------------|-----------|--------|---------------------------------------|--------------|-----------------|--|
| | OR | 95%CI | Р | OR | 95%CI | Ρ | OR | 95%CI | Р | |
| Age, years | 1.074 | 1.01-1.15 | 0.028 | | = | | | = | | |
| Preoperative MUL, mm | 0.635 | 0.53-0.74 | < .0001 | 0.699 | 0.56-0.85 | 0.0002 | 0.743 | 0.56-0.96 0. | 0.56-0.96 0.026 | |
| Console time, min | | - | | 1.007 | 0.99-1.02 | 0.167 | 0.997 | 0.95-1.05 0. | 09 | |
| Leakage at vesicourethral anastomosis | | _ | | 0.123 | 0.03-0.43 | 0.0014 | | - | | |

OR odds ratio, CI confidence interval



was associated with postoperative incontinence. They suggested that preserving MUL is important for time-to-recovery and degree of recovery. The membranous urethra contains smooth muscle fibers along its entire length and is surrounded by the rhabdosphincter [21–23]. Decreasing intraoperative trauma when preserving the MUL, which includes a greater amount of smooth muscle fibers and rhabdosphincter, has an important role in continence after RP because it contributes to maintaining and increasing urethral closure pressure [24].

Our result suggested that a longer preoperative MUL has advantages for early urinary continence recovery after RALP. In the open era, MUL might not have been taken into account as an operator's factor, but in the RALP era, the anatomical difference is evident more clearly and objectively. Therefore, not only the different operator technique, but also the MUL would relate directly to the early recovery of urinary continence.

In multivariate analysis, increasing patient age also was a risk factor for incontinence after RALP. Several studies showed a greater impact on urinary continence recovery with increasing age [25, 26]. Meanwhile, Basto et al. [27] reported urinary continence recovery rates after RALP in older men that were comparable to their younger counterparts, and thus, this should not be a reason to deny older men with a reasonable life-expectancy curative treatment for localized prostate cancer. In addition, leakage at vesicourethral anastomosis was one of the risk factors for incontinence after RALP in multivariate analysis. Leakage at vesicourethral anastomosis causes inflammatory change followed by fibrotic tissue development around the anastomotic site and sometimes causes a refractory fistula. Stavros et al. [28] reported leakage at vesicourethral anastomosis causes complications including incontinence after prostatectomy. When the MUL is < 12 mm, the patient is older or complication of leakage at vesicourethral anastomosis, early recovery of urination would not be easy. Therefore, we suggested that MUL is an important factor for preoperative informed consent regarding continence.

Our study has several limitations. First, there are some inherent limitations of pad use as an outcome measure. We defined continence as the use of one pad or less per day and did not measure pad weight. What is important in actual clinical situations is whether the number of pads is related to QOL improvement. Therefore, we investigated a QOL questionnaire for the use of a security pad. Second, MRI was performed before RALP, while MUL was measured retrospectively. In terms of measuring MUL, we evaluated MUL by a number of urologists at the preoperative conference. Therefore, the data of MUL which were summarized by a urologist who was blinded to clinical data would be more reproducible and for less selective bias in this study. Third, there was a negative impact between urinary continence and nerve sparing in our study. The aim of nerve preservation was not only for urinary continence but also for the prevention of erectile dysfunction. We also considered that cancer control was more important. Therefore, we have performed nerve preservation on one side where cancer was not detected. There was no case of bilateral nerve preservation. Although 70 cases were performed on nerve preservation, no significant difference in urinary incontinence rate between preservation group and no preservation group was shown (p = 0.312). Steineck et al. [10] reported that there is a relation between nerve preservation and urinary incontinence, but their study examined nerve preservation on both sides. In contrast, Pick

et al. [29] reported no significant difference was found in continence rates after RALP between hemi–nerve sparing and no nerve sparing. We consider that a more detailed examination about an association between nerve sparing and urinary incontinence is necessary. Finally, the retrospective design also might be a limitation. We currently are performing a prospective examination of a modified procedure on the prostatic apex according to the MUL during RALP.

Conclusion

In conclusion, preoperative MUL $> 12 \, \mathrm{mm}$ would be a predictive factor for the recovery of urinary continence at 3 months after RALP. Evaluation of preoperative MUL would be useful in clinical settings because it is easy to measure and to acquire beneficial informed consent. This result should be validated by well-conducted prospective randomized controlled trials in the future.

Abbreviations

BMI: Body mass index; EPIC: Expanded Prostate Cancer Index Composite; MRI: Magnetic resonance imaging; MUL: Membranous urethral length; PSA: Prostate-specific antigen; QOL: Quality of life; RALP: Robot-assisted laparoscopic prostatectomy; ROC: Receiver operating characteristic; RP: Radical prostatectomy; TURP: Transurethral resection of the prostate

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Availability of data and materials

The datasets supporting the conclusion of this article are included within the article.

Authors' contributions

DI contributed to the project development, data collection and management, data analysis, and manuscript drafting. YK contributed to the project development and manuscript editing. MK and RT contributed to the data analysis. AI, MO, RK, and TM contributed to the data collection. KI and WO contributed to the project development and manuscript editing. All authors read and approved the final manuscript.

Ethics approval and consent to participate

The study was approved by the institutional research committee and was performed in accordance with the ethical standards as laid down in the 1964 Declaration of Helsinki and its later amendments or comparable ethical standards. All the patients in this study gave informed consent.

Competing interests

The authors declare that they have no competing interests.

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