Comparison of Perioperative Outcomes Between Laparoscopic and Open Partial Nephrectomy for Different Complexity Renal Cell Carcinoma Based on the R.E.N.A.L. Nephrometry Score

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Purpose: This study aimed to compare perioperative outcomes of open partial nephrectomy (OPN) and laparoscopic partial nephrectomy (LPN) from a retrospective single-center dataset.

Patients and Methods: A retrospective single-center analysis of 210 patients who underwent open (n=91) or laparoscopic (n=119) partial nephrectomy for RCC was conducted between 2012 and 2015. All patients were grouped into low complexity, moderate complexity, and high complexity according to the R.E.N.A.L. Nephrometry Score, respectively. The rates of intraoperative and postoperative complications estimated blood loss, warm ischemia time, operative time, conversion of laparoscopic procedure to open surgery, and postoperative length of stay were assessed for both procedures.

Results: In low complexity group (n=93), patients undergoing LPN (n=52) under ischemia conditions had significantly longer renal artery clamp time ($p < 0.001$) and operative time ($p = 0.001$) compared with OPN (n=41). However, patients undergoing LPN had a significantly less postoperative length of stay ($p=0.005$) and estimated blood loss ($p < 0.001$) compared with OPN. There was no statistically significant difference in the rates of complications between LPN and OPN. In the moderate complexity group (n=114), 67 and 47 patients underwent LPN and OPN, respectively. LPN had notably longer warm ischemia time ($p < 0.001$) and operative time ($p < 0.001$) compared with OPN. There were no statistically significant differences in the rates of complications, estimated blood loss, and postoperative length of stay between LPN and OPN. In the high complexity group (n=3), all patients underwent OPN.

Conclusion: OPN and LPN procedures performed in patients with low and moderate complexity tumors based on the RENAL Nephrometry score offer acceptable and comparable results. When applied to low complexity tumors, our data suggest that laparoscopic NSS is an effective, minimally invasive therapeutic approach with the advantages of less blood loss, earlier hospital discharge, and more rapid convalescence.

Keywords: renal cell carcinoma, open partial nephrectomy, laparoscopic partial nephrectomy, R.E.N.A.L. Nephrometry Score

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Introduction

Over the past 2–3 decades, the incidence of kidney cancer has steadily increased all around the world.¹ The EUA Guidelines suggested that surgery is the only curative treatment for localized renal cell carcinoma (RCC).^{2,3} The partial nephrectomy

(PN) and radical nephrectomy (RN) for patients with ccRCC larger than 7 cm exhibited equivalent oncologic outcomes.⁴ Basing on quality-of-life and oncological outcomes, localized cT1 carcinomas are best treated through nephron-sparing surgery (NSS) rather than RN.^{5,6} The American Urological Association and European Association of Urology claim NSS as the operation of choice when feasible.^{7,8} The average estimated blood loss of the laparoscopic approach was universally lower open approach, but warm ischemia time (WIT) was longer.^{9,10} In a matched-pair study, the renal function was poorer in the laparoscopic PN cohort in the short-term after surgery.¹¹ There is no difference in PFS or OS between laparoscopic PN (LPN) and open PN (OPN) with laparoscopic expertise.^{12,13} NSS can be executed, either with an open or laparoscopic approach, basing on the surgeon's expertise and skills and utilizable equipment.^{14,15}

The R.E.N.A.L. Nephrometry Score can classify a tumor's complexity-grade via the anatomical characteristics of renal masses on computerized tomography/magnetic resonance imaging.^{16,17} At present, there are fewer studies on the comparison of open versus laparoscopic PN through tumor's anatomical complexity grade. The purpose of this paper is to compare perioperative outcomes of laparoscopic versus open PN adopt the R.E.N.A.L. Nephrometry Score.

Patients and Methods

Patients Collection

After institutional review board consent was achieved, we identified 210 NSS procedures executed by 5 urologic oncology surgeons at the Department of Urology of the Huashan Hospital, Shanghai, China, between June 2012 and July 2015. The inclusion criteria were as follows: (1) the pathological diagnosis was renal cell carcinoma; (2) the tumor is locally solitary; (3) patients were treated with partial nephrectomy; (4) patients with complete clinical characteristics. The surgical methods included LPN (posterior abdominal approach) and OPN (through the 11th ribs). Among the 210 patients, 93 (LPN=52, OPN=41) had 4–6 points (Low complexity group), 114 (LPN=67, OPN=47) had 7–9 points (Mediate complexity group), and only 3(OPN) had 10–12 points (High complexity group), and their medical records were retrospectively reviewed. Patient demographics (gender, age), clinical characteristics (body mass index, American Society of Anesthesiologists

[ASA] score), surgical characteristics (estimated blood loss, warm ischemia time, operative time, conversion of laparoscopic procedure to open surgery), postsurgical complications (up to 30 d after surgery), pathologic characteristics, and postoperative length of stay were compared for each technique at different levels, respectively. Complication data were retrospectively collected through view clinical records and the evaluation of complications was audited using the modified Clavien score scale.^{18,19}

Surgical Technique

LPN and OPN procedures were performed. All Minimally Invasive Partial Nephrectomy adopted a three- to the four-port method of a retroperitoneal laparoscopic approach. The open approach included variegated incisions between the 10th or 11th interspace, with most of them mini-flank incisions at the 11th rib.²⁰ Vascular control was obtained with a bulldog clamp for all partial nephrectomy patients and all of them did not undergo renal artery occlusion. Reconstruction defects were accomplished with continuous suture, and parenchymal hemostasis was accomplished with absorbable hemostatic agents in bolster configuration and matrix injection.²¹ The complications during surgical were treated with symptomatic treatment, such as timely hemostasis when bleeding and surgical intervention for urinary leakage.

Statistical Analysis

The fundamental features of the same complexity OPN and LPN groups were computed using the Fisher exact test for categorical variables and the Mann Whitney *U*-test for continuous variables via IBM SPSS Statistics v.19. All data were presented as mean $\pm$ SD, and when *P* value less than 0.05 was considered statistically significant.

Results

The Demographic Characteristics of Patients

From June 2012 to 2015, a total of 232 cases of local single RCC who underwent PN surgery with pathological diagnosis of renal carcinoma were collected. Among these patients, 210 patients (139 males and 71 females) have complete clinical characteristics were enrolled in this study. Of these 210 patients, 93 (LPN=52, OPN=41) had 4–6 points (Low complexity group), 114 (LPN = 67, OPN = 47) had 7–9 points (Mediate complexity group), and only 3(OPN) had 10–12 points (High complexity group). The mean age of the

patients was 55.7 years, range from 28 to 83 years, and the mean BMI was 21.7 kg/m^2 , range from 16.8 to 27.4 kg/m^2 . According to the ASA score, there are 138 cases of I stage, 49 cases of II stage, and 23 cases of III stage.

In the low complexity group, 52 cases received LPN, and 41 cases received OPN. And in the mediate complexity group, 67 patients received LPN and 47 patients received OPN. Additionally, in the low complexity group, the tumor radius in patients who were treated with OPN was range from 0.50 cm to 2.90 cm (mean value: 1.39 cm), and in those treated with LPN was range from 0.53 cm to 3.16 cm (mean value: 1.42 cm). In the moderate complexity group, the tumor radius of OPN was range from 0.51 cm to 2.96 cm (mean value: 1.50 cm) and LPN was range from 0.54 cm to 3.26 cm (mean value: 1.49 cm), which has no significant statistical difference. Moreover, there is no statistical difference in the R.E.N.A.L. score between LPN and OPN in the low complexity group (4 to 6 score) and moderate complexity group (7–9 score) (Figure 1). Thus, a total of 207 patients were further analyzed.

Analysis of Characteristics and Surgery Outcomes in Low Complexity Group

Firstly, the demographic and clinical characteristics of patients who received LPN or OPN were compared. The data in Table 1 revealed that the age, gender, ASA score, BMI, TNM stage, and pathological type have no significant difference (all $P > 0.05$).

Furthermore, perioperative characteristics in the low complexity group were analyzed. The estimated blood loss, operation time, renal artery clamp time, and postoperative length of stay showed a significant difference between OPN and LPN groups ($P < 0.05$, Table 1). There were no cases of intraoperative and postoperative transfusion in both LPN and

OPN groups. The results in Table 2 demonstrate that in the OPN group, estimated blood loss was about $137.1 \pm 56.1 \text{ mL}$, while in the LPN group, the estimated blood loss was about $85.6 \pm 58.5 \text{ mL}$ ($P < 0.001$). The mean duration of surgery in the OPN group was about $115.1 \pm 26.6 \text{ min}$ (55–158 min), while in the LPN group was about $142.5 \pm 51.6 \text{ min}$ (65–284 min) ($P = 0.001$). The renal artery clamp time of the OPN group was $11.9 \pm 2.6 \text{ min}$ (7–16 min), while in the LPN group was $17.0 \pm 4.2 \text{ min}$ (8–25 min). Regarding the postoperative length of stay, the average of hospitalization days was 8.0 ± 1.9 days in the OPN group and 7.0 ± 1.7 days in the LPN group ($P = 0.005$). These results revealed that patients undergoing LPN had a significantly less postoperative length of stay ($P = 0.005$) and estimated blood loss ($P < 0.001$) compared with OPN in the low complexity group.

The results in Table 1 show the postoperative complication differences of low complexity renal cortical tumors. A total of 11 cases in the OPN group had complications, 1 case of urinary leakage, and 1 case of grade III or above complications; in the LPN group, there were 7 cases with complications, 1 case of urinary leakage, and 1 case of grade III or above complications. There was no significant statistical difference between the two groups.

Comparison of Characteristics and Surgery Outcomes in Mediate Complexity Group

In the mediate complexity group, the demographic and clinical characteristics of patients who were treated with LPN or OPN have no significant difference, including age, gender, ASA score, BMI, TNM stage, and pathological type (all $P > 0.05$, Table 2). As listed in Table 2, the duration of surgery time and renal artery clamp time was 154.5 ± 48.3 and $23.4 \pm 4.8 \text{ min}$ in the LPN group,

Figure 1 The cases of patients in LPN and OPN groups. (A) The cases of patients in different scores in LPN group. (B) The cases of patients in different scores in OPN group.

Note: Anterior (a)/posterior (p).

Table 1 Perioperative and Postoperative Clinical Characteristics of Patients Treated with LPN or OPN for Low Complexity Renal Cortical Tumors

Parameters	OPN	LPN	P-value
	(n = 41) (%)	(n = 52) (%)	
Age (years)	57.9 ± 13.9	57.6 ± 9.6	0.905
Gender (male)	28(68.3)	30(57.7)	0.295
ASA score			0.947
1–2	37(90.2)	46(88.5)	
3–4	4(9.8)	6(11.5)	
BMI (kg/m ²)	22.5 ± 2.9	21.5 ± 2.7	0.867
TNM stage			0.448
pT1a	38	51	
pT1b	3	1	
Pathological type			0.200
Clear cell	35	49	
Papillary	4	3	
Chromophobe	2	0	
Estimated blood loss (mL)	137.1±56.1	85.6±58.5	< 0.001
Duration of surgery (min)	115.1±26.6	142.5±51.6	0.001
Transfusion	0	0	–
Renal artery clamp time (min)	11.9±2.6	17.0±4.2	< 0.001
Conversion to open	–	2	–
Postoperative length of stay (d)	8.0±1.9	7.0±1.7	0.005
Positive surgical margin	1(2.4)	2(3.8)	0.703
Any complication	11(26.8)	7(13.5)	0.105
Grade ≥3 complications	1(2.4)	1(1.9)	0.865
Any urine leak	1(2.4)	1(1.9)	0.865
Urine leak requiring intervention	0	0	–

Abbreviations: OPN, open partial nephrectomy; LPN, Laparoscopic partial nephrectomy; ASA, American Society of Anesthesiologists; BMI, body mass index.

respectively, while the data were 121.0 ± 26.0 and 16.2 ± 5.3 min in the OPN group. These results revealed that LPN had notably longer renal artery clamp time ($P < 0.001$) and operative time ($P < 0.001$) compared with OPN.

There were no statistically significant differences in the rates of complications, estimated blood loss, and postoperative length of stay between LPN and OPN (all $P > 0.05$, Table 2).

Discussion

The anatomy of RCC, such as diameter, depth of growth, and hilar vessel location, is considered an important predictor of surgical complexity and the occurrence of postoperative complications in partial nephrectomy for RCC.^{22,23} In 2009, Alexander Kutikov and co-workers proposed the R.E.N.A.L. scoring system based on RCC radius (R), exophytic/endophytic properties of the tumor

(E), nearness of tumor deepest portion to the collecting system or sinus (N), anterior (a)/posterior (p) descriptor (A) and the location relative to the polar line (L).¹⁶ The R.E.N.A.L. scoring system can not only accurately classify RCC into low complexity group, mediate complexity group, and high complexity group, but also predict intraoperative, postoperative outcomes and complications for patients with partial nephrectomy.^{24,25} At present, few studies are comparing laparoscopic and open partial nephrectomy for RCC based on R.E.N.A.L. score, and there is no direct objective basis for whether LPN or OPN should be chosen for patients to undergo partial nephrectomy surgery.

A previous study reported that patients treated with LPN exhibited shorter surgical time, less blood loss, shorter hospital stay, faster postoperative recovery time, but longer intraoperative warm ischemia time compared

Table 2 Perioperative and Postoperative Clinical Characteristics of Patients Treated with Partial Nephrectomy for Moderate Complexity Renal Cortical Tumors

Parameters	OPN	LPN	P-value
	(n = 47) (%)	(n = 67) (%)	
Age (years)	54.4 ± 13.7	53.8 ± 12.6	0.821
Gender (male)	33 (70.2)	53 (79.1)	0.278
ASA score			0.934
1–2	43 (91.5)	61 (91.0)	
3–4	4 (8.5)	6 (9.0)	
BMI (kg/m ²)	21.2 ± 2.5	21.9 ± 2.7	0.223
TNM stage			0.207
pT1a	41	64	
pT1b	6	3	
Pathological type			0.662
Clear cell	41	60	
Papillary	4	6	
Chromophobe	2	1	
Estimated blood loss (mL)	181.2±103.4	165.5±76.5	0.356
Duration of surgery (min)	121.0±26.0	154.5±48.3	< 0.001
Transfusion	8 (17.0)	4 (6)	0.114
Renal artery clamp time (min)	16.2±5.3	23.4±4.8	< 0.001
Conversion to open	–	5	–
Postoperative length of stay (d)	8.2±1.9	8.3±2.3	0.118
Positive surgical margin	1 (2.1)	3 (4.5)	0.529
Any complication	14 (29.8)	20 (29.8)	0.994
Grade ≥3 complications	2 (4.2)	4 (6.0)	0.686
Any urine leak	2 (4.3)	4 (6.0)	0.686
Urine leak requiring intervention	1 (2.1)	2 (3.0)	0.778

Abbreviations: OPN, open partial nephrectomy; LPN, Laparoscopic partial nephrectomy; ASA, American Society of Anesthesiologists; BMI, body mass index.

with those treated with OPN in 200 consecutive patients with a sporadic single renal tumor.²⁶ The study by Gholam Hossein Rezaeetalab and coworkers compared the difference between OPN and LPN in a group of patients over 18 years old with a single renal mass of less than 4 cm, the results indicated that LPN has some benefits over OPN, including decreased postoperative pain and higher patient satisfaction.²⁷ In the present study, the patients were divided into low complex renal tumor group, mediate complex group, and highly complex group according to the R.E.N.A.L score. All patients in the highly complex group were treated with OPN. Thus, further analyses were carried out in low and mediate complex groups, respectively. The results showed that LPN had less bleeding than the OPN group in both the low complexity group and mediate complexity group, which were generally

consistent with the results in a previous study.²⁶ These data reflected the characteristics of LPN with fine anatomy, clear vision, and less damage to the tissues and blood vessels surrounding the lesion.

The current study also observed that in terms of duration of surgery and renal artery clamp time, LPN exhibited longer time than OPN in both the low complexity group and mediate complexity group, which mainly because LPN renal reconstruction is more difficult than OPN with less space for operation and more time required for changing suture instruments, and also related to the operator's proficiency. Another single-center study also observed that the LPN group had higher warm ischemia time, intraoperative estimated blood loss, and operation duration, but exhibited shorter hospitalization time and less severe complications than the OPN group.²⁸ Whereas, both the

operative time and warm ischemia time are within an acceptable time frame, and many studies demonstrated that there is no significant difference in long-term glomerular filtration rate,^{29,30} and that operative time and warm ischemia time are not constraining factors in the choice of LPN.

In the present study, the results also exhibited that the mean postoperative hospital stay of LPN was significantly less than that of OPN in the R.E.N.A.L score low complex group. In the moderate complex R.E.N.A.L group, the mean hospital stays time after treatment of LPN was slightly longer than that of OPN, but there was no statistically significant difference. LPN has the advantage of quick postoperative recovery in RCCs with low R.E.N.A.L scores, but not in RCC patients with moderate R.E.N.A.L scores. The postoperative related complications in LPN and OPN groups in both low and moderate complex groups have no significant difference. These results were in accordance with previous studies about the results between OPN and LPN treated groups.^{29,31} A recent study by Salah M and co-workers indicated that a newly modified nephelometry score (MNS) was associated with better prediction of the outcome of partial nephrectomy when compared to R.E.N.A.L.³² In future studies, the data will try to be further compared by using the newly modified nephelometry score.

This study is a non-clinical randomized controlled retrospective case study and there are some limitations. For instance, there may be some bias in the statistics in terms of clinical data and measurement of imaging data. There is no accurate and complete data of preoperative and postoperative renal function, as well as complete data of glomerular filtration rate. Recent studies reported that off-clamp and on-clamp robotic partial nephrectomy (RPN) are safe approaches and have negligible impact on renal function.^{33,34} Furthermore, robot-assisted laparoscopic partial nephrectomy (RALPN) has been carried out in our hospital. In the future, RALPN, LPN, and OPN can be compared, and patient's glomerular filtration rate, renal function, quality of life, and survival rate can be followed up for follow-up related studies.

Conclusion

In conclusion, LPN has obvious advantages in intraoperative bleeding and postoperative recovery. There is no significant difference between LPN and OPN in postoperative related complications. RCC patients with R.E.N.A.L score (4–6) are recommended to be treated with

LPN as the first choice. R.E.N.A.L score for moderate complex renal tumors, LPN has no significant difference in postoperative complications and postoperative recovery compared with OPN and has a longer operative time and warm ischemia time. Thus, for patients with moderate complex renal tumors, the surgeon can choose the surgical approach based on his own experience and expertise. The present study provides a basis for the clinical treatment of RCC.

Ethics Statement

All procedures performed in studies involving human participants were in accordance with the ethical standards of the Shanghai Huashan Hospital research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards. Each participant signed written informed consent.

Disclosure

The authors report no conflicts of interest in this work.

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