# Application of intraoperative ultrasonography in retroperitoneal laparoscopic partial nephrectomy: A single-center experience of recent 199 cases

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# ABSTRACT

Objectives: To summarize the value and application experiences of intraoperative laparoscopic ultrasonography (ILUS) in retroperitoneal laparoscopic partial nephrectomy (RLPN). Materials and Methods: From January 2013 to December 2018, RLPN with ILUS was performed on the recent 199 patients in our center (two patients received bilateral RLPN due to suspected malignancy of both right and left sides), and the relevant clinical and follow-up data were retrospectively reviewed. Among them, 119 patients were male and 80 were female; the age of patients was  $53.4 \pm 12.3$  years. Of all the renal tumors, 105 were located on the left side and 96 on the right side with a RENAL score of  $6.6 \pm 1.7$ . All the patients were diagnosed as or suspected of having a renal tumor by preoperative imaging examination. The ILUS was applied in all the operations to help locate the tumor, delineate the boundary, clarify the diagnosis, observe the blood supply, and so on. **Results:** RLPN with ILUS in these 199 patients was successfully performed without conversion to open surgery. All surgeries were completed in 90.2  $\pm$  21.7 min, with 73.6  $\pm$  89.2 mL for estimated blood loss, and 19.3  $\pm$  5.6 min for warm ischemia time. The tumor size was  $3.6 \pm 1.5$  cm, and all the surgical margins were negative. The drainage days and postoperative hospital days were  $4.7 \pm 2.3$  and  $6.1 \pm 2.3$ , respectively. The preoperative creatinine was  $69.7 \pm 19.4 \mu$ mol/L compared with  $61.6 \pm 12.7 \mu$ mol/L measured 1 month postoperatively. There were 17 cases of renal cell carcinoma no more than 1 cm, and they were resected without artery clamp or a large amount of blood loss. Satellite tumors were confirmed in 12 cases, of which 8 were not detected by preoperative examinations and finally found by ILUS during surgeries. Conclusion: ILUS can alleviate the difficulty of preoperative diagnosis, facilitate surgical dissection, and improve the effect of nephron-sparing surgeries. Due to its great advantage, ILUS should further be promoted and applied.

Key words: Intraoperative ultrasonography, laparoscopy, partial nephrectomy, renal tumor

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## INTRODUCTION

As one of the most common malignancies in the world, kidney and renal pelvic cancer occupies the 6<sup>th</sup> and 10<sup>th</sup> leading cancers in males and females for estimated new cases, respectively. In recent years, the morbidity has increased with the advancement of medical imaging including computed tomography (CT) and ultrasound.<sup>[1]</sup> Currently, surgeries stand as the gold standard option for the renal cell carcinoma (RCC). In 1884, nephron-sparing surgery was introduced and has rapidly developed in the last 20 years.<sup>[2]</sup> For over a century, radical nephrectomy was received by most patients because nephron-sparing surgeries were confined to patients without sufficient renal function due to solitary kidney, bilateral renal tumors, or renal insufficiency.<sup>[3]</sup> With the advancement of laparoscopic techniques, retroperitoneal laparoscopic partial nephrectomy (RLPN) has been striding to treat T2 RCC, with less complications, similar oncological results, and cutting edge advantages in renal function compared to open radical nephrectomy.<sup>[2,4]</sup>

In complicated cases, anatomical anomalies, operation history, and specific tumor location could probably result in surgeries with high difficulty and potential complications. Since its first introduction in 1982, intraoperative laparoscopic ultrasonography (ILUS) has been used in urology and has gradually shown its advantage.<sup>[5]</sup> Doppler integrated into laparoscopic ultrasound allows for real-time presentation of anatomy in surgeries. Although there are articles discussing its usage in laparoscopic surgeries,<sup>[5-7]</sup> reports in RLPN are few, especially the ones dedicated in complex cases and with relative long-term and complete follow-up results.

The present study aims to describe our technique and summarize our experiences of ILUS in RLPN. We hope to provide relevant skills and experiences with ILUS to enable the urologist to better conduct puzzling and difficult retroperitoneal laparoscopic nephron-sparing surgeries.

### MATERIALS AND METHODS

## Patients

An institutional retrospective review has been approved by the ethics committee in our hospital to present the study of RLPN with ILUS. Informed consent has been obtained after necessary information was declared to the patients. RLPN was performed by the same major urologist, Nianzeng Xing, with different assistants. Perioperative data were collected independently by two researchers and discussed to be integrated into one when presented with inconsistencies. Routine hematological, biochemical examination, X-ray chest, abdominal ultrasonography, and abdominal CT/magnetic resonance imaging were assessed preoperatively.

# Surgical procedures of retroperitoneal laparoscopic partial nephrectomy with intraoperative laparoscopic ultrasonography

ILUS and its initial application in RLPN in our department have been introduced in another article published in 2012.<sup>[8]</sup> Aloka 4000 (Aloka Co., Ltd., Tokyo, Japan) ILUS machine and its matching probe (UST-5536-7.5 Ultrasound; Aloka) were applied in all cases. Surgical procedures have been standardized by the operator. After general anesthesia, ureteral stents might be placed for the tumors near the renal hilum with the guidance of cystoscopy, through which renal pelvis could be transfused with methylene blue when the suspicion of collecting system damage existed. With trocars placed and retroperitoneal space established, the surgeon opened the Gerota's fascia, and a probe could be applied to facilitate the location of the main renal artery and the accessory ones if suspicion existed. After identification of the tumor, characteristics of the tumor were compared and integrated with preoperative information, including size, location, blood supply, distance to the renal pelvis, possible pathology, and possible satellite tumors. With the confirmation of the information above, the urologist delineated the tumor with sufficient normal tissue under the guidance of ILUS and resected it after the bulldog clamp discontinued the blood flow shown by the ILUS<sup>[9]</sup> [Figure 1]. The incision was sutured with 3-0 Vicryl (Ethicon Endo-Surgery, Johnson and Johnson) in the inner plane or the renal pelvis and 2-0 Vicryl in the outer plane. The blood supply of the kidney was restored without the bulldog clamp. After confirmation of the blood supply by the ILUS, the specimen was extracted and the incisions were sealed.

The perioperative information of the recent 199 cases receiving RLPN with ILUS was retrospectively reviewed and analyzed. Preoperative information collected were age, gender, tumor features, preoperative clinical diagnosis, and so on. Operative data comprised operative time, estimated blood loss (EBL), warm ischemia time (WIT), tumor features according to the ILUS, and so on. Postoperative information included



**Figure 1.** (a) The probe of the ILUS was placed into the retroperitoneal space through the trocar and directly located on the surface of the kidney. (b) The ILUS could provide more information (tumor position, size, peripheral organs and tissue, etc.) for the surgeon. (c) The ILUS could clearly display the blood supply of the renal tumor. (d) After the renal artery was intermittently blocked, the ILUS could help us determine if the artery was completely blocked and there was accessory renal artery. ILUS: Intraoperative laparoscopic ultrasonography

complications, postoperative hospital days, the pathology of the tumors, and follow-up results.

### Statistical methods

The data were presented as mean  $\pm$  standard deviation with range for the continuous data with normal distribution and median and interquartile range for those not. One-sample Kolmogorov–Smirnov test was applied to assess whether the data are normally distributed or not. Statistical analysis was completed by Statistical Package for the Social Sciences, version 24.0 (SPSS Inc., Chicago, IL, USA).

### **RESULTS**

From 2013 to 2018, 199 patients who received RLPN with ILUS were reviewed for this study. RLPN was performed by one major urologist, Nianzeng Xing, with various assistants. Preoperative information is shown in Table 1, and operative and postoperative information is shown in Table 2.

RLPNs were all successfully completed without conversion to open surgery. All surgeries were

# Table 1. Characteristics of patients with renaltumor receiving retroperitoneal laparoscopicpartial nephrectomy

Variables	Number of variables
Patients	199
Age (mean±SD), years	53.4±12.3
BMI (mean±SD), kg/m <sup>2</sup>	25.6±3.9
Gender, <i>n</i> (%)	
Male	119 (59.8)
Female	80 (40.2)
Laterality, n (%)	
Left	105 (52.2)
Right	96 (47.8)
RENAL score, n (%)	6.6±1.7
Anterior	75 (37.3)
Posterior	65 (32.3)
Х	61 (30.4)

BMI: Body mass index, SD: Standard deviation

#### Table 2. Operative and postoperative information

Variables	Value of variables	
Operative time, min	90.2±21.7	
Warm ischemia time, min	19.3±5.6	
Estimated blood loss, mL	73.6±89.2	
Tumor size, cm	3.6±1.5	
Drainage days, days	4.7±2.3	
Postoperative hospital days, days	6.1±2.3	
Preoperative creatinine, µmol/L	69.7±19.4	
Postoperative creatinine, µmol/L (30 d)	61.6±12.7	

completed in 90.2  $\pm$  21.7 min, with 73.6  $\pm$  89.2 mL for EBL, and 19.3  $\pm$  5.6 min for WIT. The drainage days and postoperative hospital days were 4.7  $\pm$  2.3 and 6.1  $\pm$  2.3, respectively. Moreover, the preoperative creatinine was 69.7  $\pm$  19.4 µmol/L compared with 79.9  $\pm$  25.3 µmol/L on the postoperative 1<sup>st</sup> day and 61.6  $\pm$  12.7 µmol/L on the postoperative 1 month.

One patient was conversed to retroperitoneal laparoscopic radical nephrectomy because of renal pelvis invasion of the tumor shown by ILUS, which was not clearly detected by preoperative CT scan. Four patients were evaluated as Bosniak category III cysts by CT scan preoperatively. Two of them were assessed as more possibility of malignancy than benign by the ILUS intraoperatively. They were judged as clear-cell RCC by intraoperative frozen-section examination and received RLPNs thereafter. The remaining two were in the opposite situation and received renal cyst unroofing.

There was one patient whose kidney showed no blood supply by the ILUS after the sutures of the renal incision were completed. Therefore, the surgeon conducted anastomosis of the subsegmental renal

artery and main renal artery. After that procedure, the ILUS confirmed blood reperfusion and the surgery was successfully completed.

There were no complications in relation to ILUS before, during, and after RLPN. Fifteen (7.54%) patients suffered from postoperative complications in this study, with abdominal pain in three patients, fever in six patients, urinary infection in four patients, and pulmonary infection in two patients. All these complications were treated well with conservative or anti-inflammatory therapy. There were no long-term complications (>90 days) present [Table 3].

The pathological reports showed that there were 149 cases of clear-cell RCC, 9 cases of chromophobe RCC, 7 cases of papillary RCC, 4 cases of oncocytoma, 23 cases of angiomyolipoma (AML), 2 cases of multilocular cystic RCC, 3 cases of renal cyst, 1 case of inflammatory lesion, 1 case of adenofibroma, and 2 cases of Xp11.2 translocation. Of all the clear-cell RCC cases, there were 15 cases of Grade 1, 118 cases of Grade 2, and 16 for Grade 3. As for the papillary RCC, there were 5 cases of Type 1 and 2 cases of Type 2. In all 169 malignant pathological results, 119 cases belonged to pT1a, 49 cases belonged to pT1b, and 1 case belonged to pT2a. Detailed information is shown in Table 4. Twelve cases of satellite tumors were confirmed by pathological results, of which 8 were not detected by preoperative examinations and finally found by ILUS during surgeries. As reported in this study, there were 17 cases of RCC no more than 1 cm, and they were resected without artery clamp or a large amount of blood loss.

During the median postoperative follow-up of 42 months, the relapse-free survival rate and the overall survival rate were 99.0% and 100%, respectively. Of the 199 patients, a total of 15 patients were lost to follow-up. Two patients relapsed postoperatively during which one patient was pathologically diagnosed with clear-cell carcinoma and lung metastasis and received target therapy, while the other patient was pathologically diagnosed with AML and received regular observation.

### DISCUSSION

Between 2013 and 2018, ILUS was applied in RLPNs and relative experiences were summarized. In this article, the authors described the experiences and its advantages in various situations in partial nephrectomy.

# Table 3. Postoperative complications and treatment

Clavien grade	n	Treatment
1	3	Conservative
1	6	Conservative
1	4	Antibiotics
1	2	Antibiotics
	0	
	Clavien grade 1 1 1 1	Clavien grade n   1 3   1 6   1 4   1 2   0

# Table 4. Postoperative pathology, T stage andgrading of renal tumors

Variables	Number of variables		
Pathology, n	201*		
Clear-cell RCC	149		
Chromophobe RCC	9		
Papillary RCC	7		
Multilocular cystic RCC	2		
Xp11.2 translocation	2		
Angiomyolipoma	23		
Oncocytoma	4		
Renal cyst	3		
Adenofibroma	1		
Inflammatory lesion	1		
T stage (pT), n			
1a	119		
1b	49		
2a	1		
Grading of clear-cell RCC, n			
Grade 1	15		
Grade 2	118		
Grade 3	16		

\*Two patients received bilateral RLPN and there are totally 201 pathological results. RLPN: Retroperitoneal laparoscopic partial nephrectomy, RCC: Renal cell carcinoma

Combined with preoperative CT and preoperative ultrasonography, ILUS can present additional real-time information to recuperate the surgeons' tactile feedback about the size, nature, number, peripheral organs and tissue, and location of the tumor in certain cases. Moreover, ILUS can provide higher resolution of the image that surgeons focused on than that of the standard transcutaneous abdominal ultrasonography as well as CT.

As an indispensable procedure, renal artery clamp is essential to the success of RLPN. Accessory renal arteries exist in certain population, and preoperative CT cannot identify each one of the arteries.<sup>[10]</sup> Moreover, preoperative CT cannot precisely indicate the location of the arteries, especially after manipulation and dissection in surgeries. In addition, various anomalies, different position during surgeries and CT, and various tumor characteristics may pave a rough road to RLPN. Once injuries of blood vessels occur, major complications or conversion is inevitable to the surgeons, anesthetists, and patients who are confronted with it. To tackle this dilemma, tentative surgical techniques or ILUS is badly needed for vascular control in RLPNs, which helps shorten operative time and provides high resolution of anatomy to the surgeons.

In general, only after occlusion of the main renal arteries, RLPN can be implemented by the surgeons. In certain cases, highly selective renal blood control is an inevitable step to achieve surgical success. Some articles have declared that highly selective artery control could result in advanced renal function compared to without it.<sup>[11,12]</sup> In this study, when segmental renal arteries were required, ILUS, an essential tool, was a remover of obstacles and clearly showed the segmental renal artery that the surgeons needed.

Although preoperative CT usually showed tumor characteristics, such as tumor boundary, ILUS still played a key role in delineating the boundary between the tumor and the surrounding tissue, especially during surgical movements. In RLPNs, posterior tumors were relatively easier to be resected than the anterior ones. ILUS was absolutely necessary for certain circumstances as follows. While RLPNs for anterior tumors must be implemented just as for posterior ones, reposition of the kidney inevitably lowers the applicability of the information provided by preoperative imaging. For tumors near the renal hilum and internal renal tumors with intricate boundary, local anatomy might be altered by pneumoperitoneum and collapsed collecting system after renal artery control.<sup>[13]</sup> Patients with operation history, especially urinary ones, were associated with complex surgical steps to tackle disordered local tissue cautiously. In these complicated situations, ILUS could be used repeatedly when the surgeon was irresolute of the local anatomy. Moreover, potential complications could be reduced when ILUS was used in circumstances as narrated above.<sup>[14]</sup> Small and flexible probe of ILUS could detect an overwhelming majority of complicated structures during the operation. The preoperative surgical plan could be changed accordingly due to this technique. In this study, no complications were induced by this technique.

As was reported previously, satellite lesions exist in renal tumors. This center had published that there were no satellites in that study.<sup>[8]</sup> However, in the following cases, 12 cases of satellite tumors have been confirmed by pathological results. It was a remarkable fact that 8 out of 12 cases had not been detected by preoperative examinations and finally found by ILUS during surgeries. To remove tumors with 5–10 mm of renal parenchyma, satellite lesions should be detected and excised by the urologists. For this reason, not only renal tumors were routinely inspected by ILUS, but also the rest of the kidneys were scanned in this study.

Malignant tissue including renal tumors and satellite lesions should be resected integrally with negative margin. On the other hand, the more renal parenchyma that was preserved, the more renal function that was probably retained.<sup>[15,16]</sup> Without ILUS, urologists have to excise more tissue than sufficient to ensure a negative pathological result. While ILUS was applied to define the surgical location, and figure out whether there was invasion to the collecting system, and satellite lesions around the tumor, especially for intraparenchymal tumors and the ones near renal hilum,<sup>[17-19]</sup> fine surgical movement and surgical plan alteration may still come into reality.

Preoperative medical imaging could not make a definite diagnosis in certain renal tumors. Owing to relatively low spatial resolution of CT and indirect transduction of ultrasound, extra information is badly needed in surgeries without definite diagnoses and the affirmative operative procedures. As shown in Figure 2, the ILUS scanned a lesion inside the kidney and indicated the possibility of malignancy. However, the preoperative CT



**Figure 2.** The ILUS scanned a lesion inside the kidney and indicated the possibility of malignancy, which the preoperative CT only provided a vague imaging evidence to the clinicians and brought great difficulty for the accurate diagnosis. ILUS: Intraoperative laparoscopic ultrasonography; CT: Computed tomography

provided only a vague imaging evidence to the clinicians and brought greater difficulty to us for the accurate diagnosis. As a result, we determined to implement RLPN and completely resected the tumor with the help of ILUS. Postoperative pathological results showed that the tumor was a renal cyst combined with clear-cell RCC, Grade 1.

Renal tumors no more than 4 cm are classified as small RCC.<sup>[20,21]</sup> With the development of medical imaging, 2 cm might also be kept as a threshold of small renal lesion according to our experience. For lesions no more than 2 cm, renal artery clamp might not be necessary, but the tumors are intricate and must be located and resected precisely during surgeries. To eradicate malignant tissues and protect the renal parenchyma, perirenal fat should not be peeled off especially close to the tumor, which facilitates manipulation of the tumor and the kidney. Preoperative CT usually indicates tumor characteristics including location, but after dissection, vague location might become the reality that surgeons are confronted with.<sup>[22,23]</sup> In this study, there are 17 cases of RCC no more than 1 cm, and they were resected without artery clamp or a large amount of blood loss. ILUS was essential and necessary in these 17 surgeries for its convenience, high accuracy, and low cost. When it comes to functionally solitary kidney or isolated kidney, RLPN may pose a more severe position to the patients and doctors for its relatively poor function. ILUS should be applied in case that negative margin dominates the surgical movement to resect more than necessary parenchyma.<sup>[24]</sup>

After sutures of the renal incisions, the ILUS should be applied to detect the renal blood supply in the medical center for an event. As reported above, the renal blood supply of one patient could not be detected. The reason why there was no blood reperfusion is still unknown to the surgeons, but the view that ILUS should be applied with full-scale necessity after sutures is engraved to the mind of clinicians in this center although the incidence of similar events is sheer low. ILUS could show significant advantages for the assessment of tissue and vascular control for abdominal organs.<sup>[25]</sup> In this cohort, no positive surgical margin has been reported with the facilitation of ILUS to some certain degree.

# CONCLUSION

In summary, with ILUS demarcating the boundary of the tumor simultaneously and careful practice, negative surgical margins can be achieved by experienced urologists, so that more renal parenchyma can be preserved for the patient. In general, ILUS is a vital and convenient tool to guide RLPN in multiple respects with relatively high resolution and rare concomitant complications.

There are several limitations existing in this article for its retrospective, single-surgeon, and single-center study design. Moreover, surgical experience was a subjective factor that cannot be neglected and quantified. However, due to the large quantity of sample size, comprehensive description of relatively rare cases, and long-term follow-up, this study exhibited the particular experience and surgical application of ILUS to promote further development of this technique.

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### Conflicts of interest

There are no conflicts of interest.

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