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## Reoperative Partial Nephrectomy—Does Previous Surgical Footprint Impact Outcomes?

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**Abstract**

**Purpose:** Historically, open techniques have been favored over minimally invasive approaches for complex surgeries. We aimed to identify differences in perioperative outcomes, surgical footprints, and complication rates in patients undergoing either open or robotic reoperative partial nephrectomy.

**Materials and Methods:** A retrospective review of patients undergoing reoperative partial nephrectomy was performed. Patients were assigned to cohorts based on current and prior surgical approaches: open after open, open after minimally invasive surgery, robotic after open, and robotic after minimally invasive surgery cohorts. Perioperative outcomes were compared among cohorts. Factors contributing to complications were assessed.

**Results:** A total of 192 patients underwent reoperative partial nephrectomy, including 103 in the open after open, 10 in the open after minimally invasive surgery, 47 in the robotic after open, and 32 in the robotic after minimally invasive surgery cohorts. The overall and major complication (grade 3) rates were 65% and 19%, respectively. The number of blood transfusions, overall complications, and major complications were significantly lower in robotic compared to open surgical cohorts. On multivariate analysis, the robotic approach was protective against major complications (OR 0.3,  $p = 0.02$ ) and estimated blood loss was predictive (OR 1.03,  $p = 0.004$ ). Prior surgical approach was not predictive for major complications.

**Conclusions:** Reoperative partial nephrectomy is feasible using both open and robotic approaches. While the robotic approach was independently associated with fewer major complications, prior approach was not, implying that prior surgical approaches are less important to perioperative outcomes and in contributing to the overall surgical footprint.

**Keywords**

robotics; minimally invasive surgical procedures; carcinoma; renal cell; von Hippel-Lindau disease

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Partial nephrectomy (PN) remains the preferred treatment for patients with small renal masses and for patients at risk for recurrent renal cell carcinoma (RCC) due to hereditary forms of kidney cancer.<sup>1</sup> Minimally invasive surgery (MIS) has shown improved length of stay and estimated blood loss (EBL) compared to open surgery while maintaining

equivalent oncologic efficacy.<sup>2</sup> However, for patients who have undergone prior partial nephrectomies, the obliteration of surgical planes and subsequent adhesions can create a hostile surgical field. Traditionally, surgeons have opted for an open surgical approach when facing more complex operations; however, there are few data to support the use of open or MIS techniques over the other in this scenario.

High quality evidence of outcomes in reoperative renal surgery is lacking, with most data being limited to small case series. Furthermore, the use of MIS rather than open surgery in these complex reoperations has an even greater dearth of information. Reoperative partial nephrectomy (RePN) via open approach was shown to be feasible; however, initial reports demonstrated a high rate of complication of 20% to 50% in all patients and in those with solitary kidneys, respectively.<sup>3,4</sup> However, as experience with this operation grew, surgeons were able to obtain similar perioperative outcomes and complication rates when matching reoperative surgical outcomes to index surgical outcomes.<sup>5</sup> Outcomes of laparoscopic PN in reoperative surgery, however, have been mixed with some series showing conversion rates to open approach as high as 40% while others demonstrated similar outcomes to procedures performed in nonoperated fields.<sup>6,7</sup> Fortunately with increasing experience and the incorporation of the robotic surgical platform, outcomes comparable to PN in nonoperated fields have been reported, though most of these series are limited by decidedly small sample sizes of fewer than 30 patients.<sup>8,9</sup>

It is also unclear if the operative approach, open or MIS, utilized in the index surgery alters subsequent operations on that renal unit. This concept of a “surgical footprint,” in which a surgical intervention in a particular operative field can affect future surgeries, may play a role in reoperative outcomes. Evidence from other surgical specialties has shown that a larger surgical footprint (ie more complete dissection, presence of an anastomosis) has led to increased postoperative complications.<sup>10</sup> Currently it is unclear if the surgical footprint differs between open and MIS surgery as the data are inconsistent across studies.<sup>10–12</sup> However, the preponderance of evidence suggests that a higher number of reoperations on a kidney leads to increased complications. Consequently, to minimize the likelihood of complications, a better understanding of the impact of the surgical footprint is needed.<sup>13</sup>

The primary goal of this study is to identify any differences in perioperative outcomes between the robotic and open approach to RePN. The next goal is to assess if the prior surgical approach caused a distinct surgical footprint, causing differences in outcomes. The final aim of this study is to identify which factors lead to increased numbers of significant complications. We hypothesize that MIS outcomes would be noninferior to open outcomes in patients undergoing RePN and that a prior MIS approach would contribute to a smaller surgical footprint compared to prior open surgery.

## MATERIALS AND METHODS

An Institutional Review Board-approved prospectively maintained registry was queried for all patients from January 2010 to July 2019 who underwent extirpative renal surgery after having prior surgery on the ipsilateral kidney (IRB No. NCI-97-C-0147). Prior surgeries were defined as minimally invasive if they were laparoscopic or robotic PN. Patients’ prior

surgical histories were reviewed, and patients were subsequently stratified into 4 cohorts: open after open surgery, open after MIS only, robotic after open surgery, robotic after MIS only (fig. 1). The cohorts are defined by the surgical approach utilized in prior renal surgery and the surgical approach used for the surgery currently being evaluated. Any history of prior open PN would automatically place the patient in a prior open surgery cohort.

Patient information was extracted including demographic, perioperative, and pathological outcomes. Complications were defined utilizing the Clavien-Dindo classification schema.<sup>14</sup> A single aspect of the classification system was modified due to the fact that all patients undergoing renal surgery at our institution are routinely monitored in the intensive care unit (ICU) until postoperative day 1. This ICU stay was not considered a complication unless patients required ICU level care due to life-threatening complication as addressed by the Clavien-Dindo schema. Glomerular filtration rate (GFR) was estimated using the Modification of Diet in Renal Disease equation. Associations between cohorts, demographic data, perioperative data, and outcomes were evaluated with the use of univariate and multivariate linear and logistic regression, 1-way analysis of variance, and Kruskal-Wallis H test for the 4-cohort analysis. Similar analysis was performed utilizing Pearson's chisquared test for categorical variables or Student's t-test and Wilcoxon-Mann-Whitney test for continuous variables for the remaining analysis. Statistical analysis was performed with STATA 16 (StataCorp, College Station, Texas). A p value of <0.05 was considered significant.

## RESULTS

From January 2010 to July 2019, a total of 672 patients underwent PN at our institution. Of these, 192 patients who had prior partial nephrectomies were identified and included in the final analysis (fig. 1). Demographic data were similar among all cohorts with only tumor laterality differing among the groups (table 1). The cohorts were divided as follows: 103 patients were in the open after open cohort, 10 were in the open after MIS only cohort, 47 were in the robotic after open cohort, and 32 were in the robotic after MIS only cohort. The proportion of robotic procedures increased during the study period. While only 24% (24 of 98) of procedures in the first half of the study period were performed via robotic approach, 59% (55 of 94) were performed via a robotic approach in the second half. A significant notable difference was the open after open cohort having an increased number of prior mean procedures at 2.0 ( $p < 0.001$ ). The majority of patients had a known hereditary cancer predisposition syndrome. A total of 124 patients (65%) experienced a complication while 36 patients (19%) experienced a grade 3 complication.

Intraoperative outcomes demonstrated notable differences among the cohorts (table 2). Units of packed red blood cells (pRBCs) transfused intraoperatively ( $p = 0.03$ ) were significantly lower in both robotic surgery arms. The median duration of operation was 388 minutes with no considerable difference noted among the cohorts. There was, however, a significant difference in the number of tumors resected at time of operation. The overall median number of tumors resected per procedure was 5, with both open surgical cohorts having significantly more tumors resected ( $p = 0.01$ ). One patient (0.5%) in the robotic after open cohort

required a conversion to radical nephrectomy while 6 patients (7.6%) in the robotic cohorts required conversion to open surgery.

Postoperatively, the number of pRBC transfusions ( $p < 0.001$ ), mean number of grade 3 complications ( $p = 0.046$ ), and number of patients experiencing any complication ( $p < 0.001$ ) were significantly lower in both robotic surgery cohorts compared to the open surgical cohorts. The number of patients who experienced a grade 3 complication was significantly lower in only the robotic after open cohort (OR 0.3,  $p = 0.02$ , table 3). There were no differences in change in 3-month median change in estimated GFR ( $p = 0.4$ ).

When assessing for factors associated with a patient developing a grade 3 complication, several demographic and intraoperative variables were found to be significant. Surgery duration (OR 1.005, 95% CI 1.001–1.008,  $p = 0.01$ ), surgical approach utilized (OR 0.3, 95% CI 0.1–0.8,  $p = 0.02$ ), and number of prior open procedures (OR 1.6, 95% CI 1.1–2.4,  $p = 0.02$ ) were significant risk factors (table 3 and fig. 2). For perioperative factors, EBL (OR 1.0004, 95% CI 1.0002–1.0005,  $p < 0.001$ ), pRBCs transfused intra and postoperatively (OR 1.2, 95% CI 1.1–1.3,  $p < 0.001$  and OR 1.5, 95% CI 1.2e–1.8,  $p < 0.001$ , respectively), and the percent change in 3-month GFR were also associated with the presence of grade 3 complication (OR 0.97, 95% CI 0.95e0.99,  $p = 0.02$ ).

Multivariate models demonstrated that EBL (OR per 100 ml: 1.03, 95% CI 1.009–1.05;  $p = 0.005$ ) was predictive of which patients experienced a grade 3 complication while being part of the robotic after open cohort was protective (OR 0.2, 95% CI 0.07–0.9,  $p = 0.03$ ) (supplementary table 1, <https://www.jurology.com>). The model was subsequently adjusted, and the 4-cohort design was replaced with the current surgical approach utilized and a history of open surgery as separate variables. Similarly, EBL (OR per 100 ml: 1.03, 95% CI 1.009–1.05;  $p = 0.004$ ) was predictive of grade 3 complications while currently undergoing robotic surgery was protective (OR 0.3, 95% CI 0.1–0.8,  $p = 0.02$ ; table 4). Notably a history of open procedures (OR 0.8,  $p = 0.7$ ) and the number of tumors resected (OR 1.0,  $p = 0.4$ ) were not associated.

Given that some patients had both prior open and MIS surgery, a subgroup analysis was performed in patients who had a history of only open PN or a history of only prior MIS PN. On multivariate analysis, only EBL was found to be predictive of a grade 3 complication (OR per 100 ml: 1.03, 95% CI 1.008–1.06,  $p = 0.007$ , supplementary table 2, <https://www.jurology.com>).

## DISCUSSION

RePN is a challenging surgery. Normal surgical planes can become altered or altogether obliterated and peri-hilar fibrosis can make vascular dissection a hazardous ordeal (fig. 3). This study is the largest to comprehensively describe the outcomes of patients undergoing complex RePN. Most notably, these results demonstrate that in experienced hands, RePNs are both feasible and safe. Of the 192 RePNs performed, only 1 patient (0.5%) required a radical nephrectomy, which was necessitated due to a vascular injury. Of the 79 attempted robotic RePN, 6 (7.6%) required conversion to open approach. These results help clarify

that repeat surgery, independent of the approach, is feasible without compromising the final outcome of a successful operation. Current literature suggests that robotic RePN is likely feasible; however, with fewer than 40 patients reported in the literature, power to make meaningful conclusions is lacking. The favorable results of the 79 patients in our robotic cohorts clearly demonstrates the feasibility of robotic RePN.

The results among the cohorts are split by current surgical approach, with both robotic cohorts experiencing decreased need for intra and postoperative transfusions, fewer patients experiencing a complication, and lower number of grade 3 complications experienced per patient and only the robotic after open cohort experiencing fewer patients with a grade 3 complication. On both multivariate models, 2 major risk factors were identified as significant contributors to experiencing a grade 3 complication: EBL and current robotic surgery. The subgroup analysis demonstrated that a history of open only or MIS only surgery was not a factor in predicting the presence of a grade 3 complication. These results suggest that the prior surgical approaches are less important in determining perioperative outcomes, and therefore less important in the overall surgical footprint. Likely it is the extent of tissue plane manipulation or dissection, whether open or robotic, that plays a greater role in the surgical footprint as is echoed in prior literature.<sup>10,15</sup>

EBL as a factor contributing to increased high grade complications is logical, as many times these patients require considerable intraoperative blood products and fluid resuscitation leading to ICU level care due to longer times weaning ventilation and other secondary issues such as pre-renal acute kidney injury. However, it is unclear if EBL is merely a predictor of the complexity of the operation and may reflect unaccounted for variables such as tumor complexity or perirenal fibrosis. Unfortunately, no validated renal mass complexity scoring system exists for patients with multifocal tumors and therefore tumor complexity could not be quantified.<sup>16</sup>

It is important to note that while the size of the surgical footprint vis-à-vis open versus minimally invasive procedures did not seem to impact outcomes, the surgical technique utilized by surgeons at our institution incorporates several factors in minimizing the footprint of surgery regardless of approach. For instance, Gerota's fascia is preserved and reapproximated at the end of each case when technically feasible. Additionally, hilar dissection is minimized, and the renal vein and artery are often not skeletonized in order to potentially limit fibrosis during subsequent procedures. Similarly, the hilum is not clamped in the majority of cases and renovascular occlusion is reserved for deeper tumors or those located near the hilum.

This study has its limitations. The study is retrospective and comprised mostly of a relatively uncommon group of patients with known or suspected hereditary renal cell carcinoma; however, the challenges and complications arising from prior surgery are not different from those with sporadic kidney cancer. There is also likely a considerable selection bias in choosing operative approach, and indeed, the open after open RePN cohort had the most prior procedures. The vast majority of patients underwent tumor enucleations in order to spare as much parenchyma as this benefits a population which is afflicted with multiple, recurrent masses. Also, the patients' prior surgeries consisted of a heterogeneous mix of

approaches with many having had both MIS or open surgery in the past. Furthermore, some patients may have had a history of focally ablative procedures. In order to standardize this group, we hypothesized that open surgery would have the most manipulation of surgical planes and therefore our definition of prior open surgery was a patient who ever had a prior open PN in the ipsilateral renal unit. The purpose of the subgroup analysis was to address this limitation by allowing for a purer assessment of the effects of prior surgical approaches, but it came at the cost of statistical power as the patient cohort size was considerably reduced. Therefore, the favorable surgical outcomes in robotic surgery do not necessarily tout robotic assisted surgery as a safer approach, but rather demonstrate its safety and feasibility in the reoperative setting. However, even with its drawbacks, this study remains the largest study analyzing patients undergoing RePN and their outcomes.

## CONCLUSIONS

In patients with a history of prior ipsilateral PN, the use of robotic approach to PN is safe, feasible, and may have improved perioperative outcomes over open surgery. The surgical footprint of prior surgeries is likely less due to the surgical approach utilized, but rather the amount of tissue plane manipulation and dissection that was performed at that time.

## Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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## Abbreviations and Acronyms

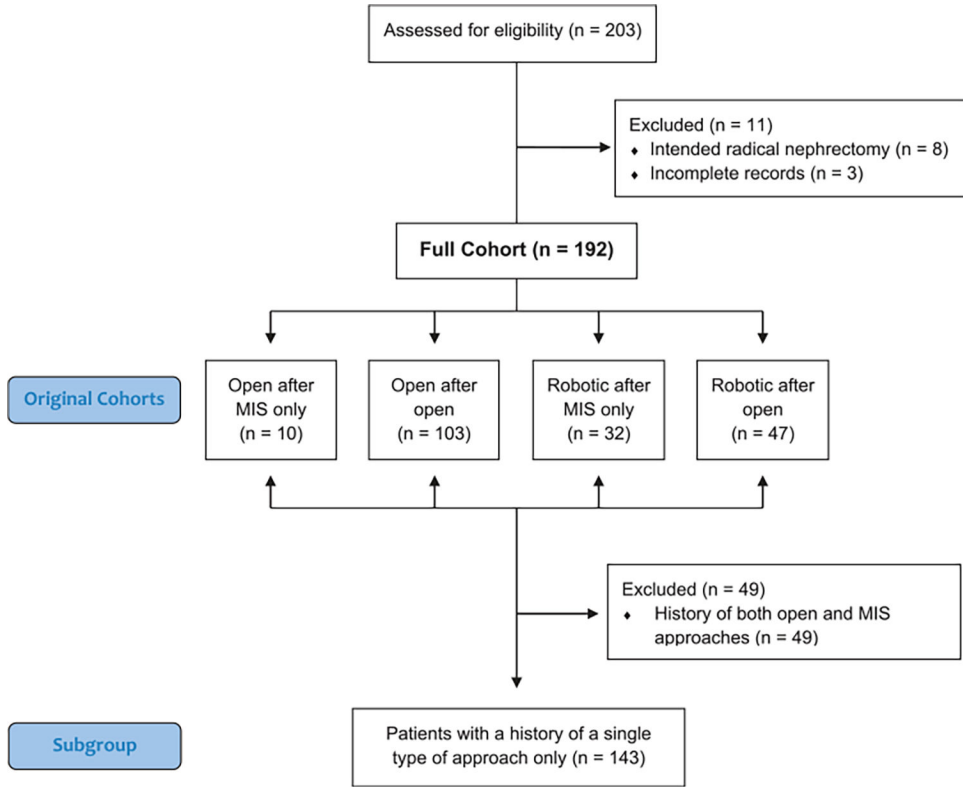
<b>EBL</b>	estimated blood loss
<b>GFR</b>	glomerular filtration rate
<b>ICU</b>	intensive care unit
<b>MIS</b>	minimally invasive surgery
<b>PN</b>	partial nephrectomy
<b>pRBC</b>	packed red blood cell
<b>RCC</b>	renal cell carcinoma
<b>RePN</b>	reoperative partial nephrectomy

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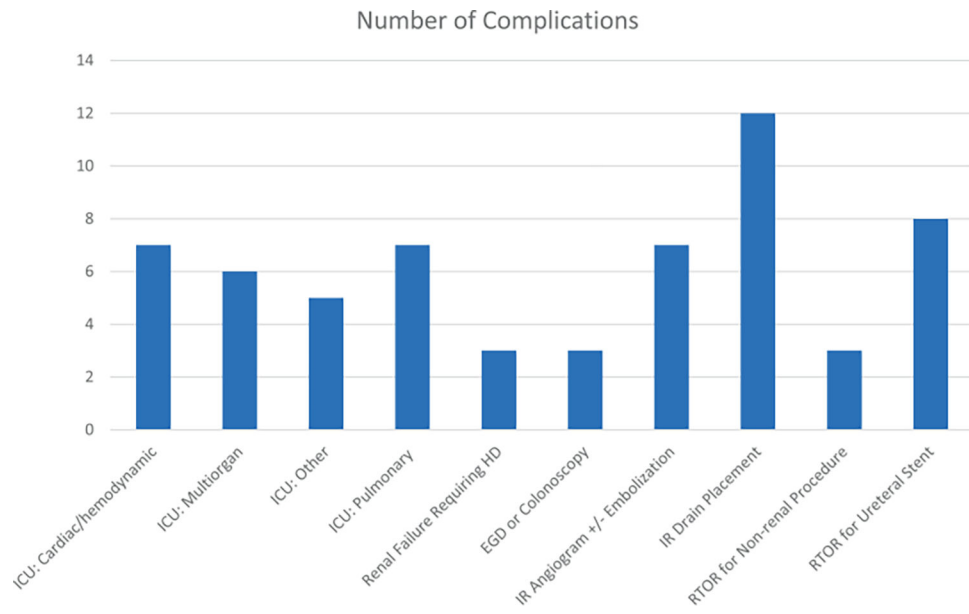
**Figure 1.** Patient cohorts. All patients were originally assigned to cohorts based on prior surgical history and current surgical approach. Any history of open surgery would place patient in prior open cohort. To remove confounding factors when assessing role of prior surgery’s effect on surgical footprints, new separate groups were created. Given that some patients had both prior open and MIS surgery, subgroup analysis was performed in patients who had history of only open partial nephrectomies or history of only prior minimally invasive partial nephrectomies.

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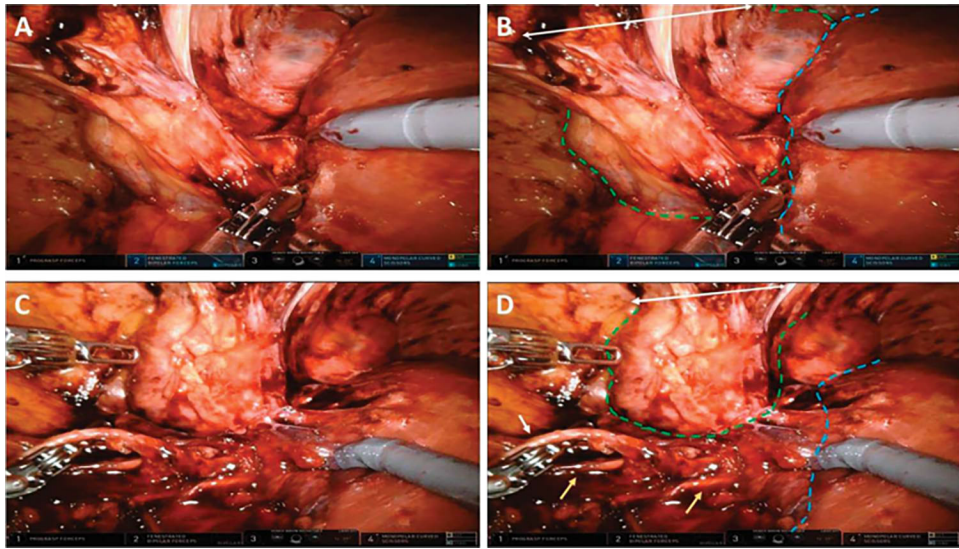
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**Figure 2.** Grade 3 complications by cause. *HD*, hemodialysis. *EGD*, esophagogastroduodenoscopy. *IR*, interventional radiology. *RTOR*, return to operating room.



**Figure 3.** Intraoperative view of robotic after open PN demonstrating characteristic peri and paranephric adhesions to anterior abdominal wall and surrounding organs and tissue. *A* and *B*, right kidney (outlined in green dashes) adherent to liver (outlined in blue dashes) and adherent to prior incision site on anterior abdominal wall (white double arrow). *C* and *D*, same patient after further dissection. White arrow indicates ureter, yellow arrow indicates right gonadal veins.

Table 1.

Patient demographics

	All Pts	Open after MIS Only	Open after Open	Robotic after MIS Only	Robotic after Open	p Value
No. pts	192	103	47	46	0.5	
Median yrs age (range)	48 (27-79)	50 (27-77)	47 (27-69)	46 (28-68)	0.9	
No. sex (%):						
Male	123 (64)	68 (66)	21 (66)	28 (60)		
Female	69 (36)	35 (34)	11 (34)	19 (40)		
No. race (%):					0.6	
White, nonHispanic	152 (79)	78 (76)	28 (88)	39 (83)		
Black/African American	116	77	13	36		
Asian	95	77	0	24		
Other	2,010	1,111	39	36		
No. hereditary predisposition syndrome (%):					0.2	
von Hippel Lindau	156 (81)	88 (85)	22 (69)	39 (83)		
Birt-Hogg-Dubé	6 (3)	3 (3)	2 (6)	1 (2)		
Hereditary leiomyomatosis + RCC	4 (2)	2 (2)	0 (0)	1 (2)		
Other	23 (12)	9 (9)	6 (19)	6 (13)		
Sporadic	2 (1)	1 (1)	2 (6)	0 (0)		
No. predominant tumor histology (%):					0.3	
Clear cell	168 (88)	94 (91)	28 (88)	40 (85)		
Papillary type 1	7 (4)	2 (2)	2 (6)	2 (4)		
Hybrid oncocytic/chromophobe tumor	6 (3)	3 (3)	2 (6)	1 (2)		
Hereditary leiomyomatosis + RCC-associated RCC	4 (2)	2 (2)	0 (0)	1 (2)		
Other	6 (3)	2 (2)	0 (0)	3 (6)		
No. American Society of Anesthesiologists® score (%):					0.994	
1	0 (0)	0 (0)	0 (0)	0 (0)		
2	1 (0.5)	0 (0)	1 (3)	0 (0)		
3	189 (98)	103 (100)	29 (91)	47 (100)		
4	2 (1)	0 (0)	2 (6)	0 (0)		
No. laterality (%):					0.03	

	All Pts	Open after MIS Only	Open after Open	Robotic after MIS Only	Robotic after Open	p Value
Lt	94 (49)	5 (50)	48 (47)	23 (72)	18 (38)	
Rt	98 (51)	5 (50)	55 (53)	9 (28)	29 (62)	
Mean±SD	1.7±0.9	1.3±0.5	2±0.9	1.3±0.8	1.5±0.7	0.0001
Median No. prior ipsilat procedures (IQR)	1 (1-2)	1 (1-2)	2 (1-2)	1 (1-1)	1 (1-2)	0.0001
No. history of ablation (%):						0.6
No	157 (82)	8 (80)	81 (79)	28 (88)	40 (85)	
Yes	35 (18)	2 (20)	22 (21)	4 (13)	7 (15)	

**Table 2.**

Perioperative factors and outcomes by cohort

	All Pts	Open after MIS Only	Open after Open	Robotic after MIS Only	Robotic after Open	p Value
No. pts	192	10	103	32	47	
Intraop characteristics and outcomes:						
No. radical nephrectomy conversions (%)	1 (0.5)	0 (0)	0 (0)	0 (0)	1 (2)	0.5
No. robotic to open conversions (%)	6 (3.1)	Not applicable	Not applicable	2 (6.3)	4 (8.5)	0.5
Median mins duration of surgery (IQR)	388 (339–460)	414 (256–475)	380 (333–432)	421 (356–472)	384 (338–489)	0.5
Median ml EBL (IQR)	2,000 (1,100–3,500)	2,100 (850–3,500)	2,100 (1,200–4,000)	1,750 (750–3,000)	1,900 (800–2,750)	0.1
Median units intraop pRBCs transfused (IQR)	3 (0–6)	2 (1–3)	41 (1–8)	2 (0–4)	2 (0–4)	0.03
No. hilar clamping utilized (%)	42 (22)	5 (50)	21 (20)	5 (16)	11 (23)	0.2
Median No. tumors resected (IQR)	5 (3–10)	6 (1–15)	7 (4–12)	4 (2–6)	4 (2–8)	0.01
Postop outcomes:						
Median units postop pRBCs transfused (IQR)	0 (0–2)	1 (1–2)	2 (0–2)	0 (0–1)	0 (0–0)	<0.001
No. pts experiencing any grade Clavien complication (%)	124 (65)	8 (65)	83 (81)	17 (53)	16 (34)	<0.001
No. pts experiencing grade 3 Clavien complication (%)	36 (19)	2 (20)	26 (25)	4 (13)	4 (9)	0.07
Mean±SD No. grade 3 Clavien complications	0.3±0.8	0.2±0.4	0.5±1.0	0.1±0.3	0.08±0.3	0.046
Median % 3-mos change in glomerular filtration rate (IQR)	-7.2 (-17.1–3.5)	-2.8 (-18.1–11.1)	-6.7 (-19.1–4.1)	-6.1 (-13.1–4.4)	-8.5 (-17.1–2.1)	0.7

**Table 3.** Univariate analysis of perioperative factors and outcomes by presence of grade 3 complication

	No Grade 3 Complications	Grade 3 Complication	Odds Ratio (95% CI)	p Value
No. pts	156	36		
Demographic:				
Median yrs age (range)	48 (27–79)	51 (28–69)	1.01 (1.0–1.04)	0.5
No. von Hippel Lindau diagnosis (%)	126 (81)	30 (83)	0.8 (0.3–2.2)	0.7
Median No. prior ipsilateral procedures (IQR)	1 (1–2)	2 (1–2)	1.3 (0.9–1.9)	0.2
No. cohort (%):				
Open after MIS only	8 (5)	2 (6)	0.7 (0.2–3.7)	0.7
Open after open	77 (49)	26 (72)	Reference	Reference
Robotic after MIS only	28 (18)	4 (11)	0.4 (0.1–1.3)	0.1
Robotic after open	43 (28)	4 (11)	0.3 (0.1–0.8)	0.02
Intraop characteristics and outcomes:				
Median mins duration surgery (IQR)	379 (325–441)	445 (373–493)	1.005 (1.001–1.008)	0.01
Median ml EBL (IQR)	1,850 (1,000–3,000)	3,500 (1,600–5,000)	1.0004 (1.0002–1.0005)	<0.001
No. hilar clamping utilized (%)	37 (24)	5 (14)	0.5 (0.2–1.4)	0.2
Median No. tumors resected (IQR)	5 (2–9)	8 (5–13)	1.04 (1.0002–1.09)	0.07

**Table 4.**

Multivariate analysis of risk factors for grade 3 complication

<b>Risk Factor</b>	<b>Logistic Coefficient</b>	<b>Odds Ratio</b>	<b>95% CI</b>	<b>p Value</b>
Surgery duration	0.003	1.003	0.998–1.008	0.2
Current surgical approach: robotic	–1.3	0.3	0.1–0.8	0.02
History of prior open surgery	–0.2	0.8	0.3–2.4	0.7
No. prior ipsilat procedures	–0.04	1.0	0.6–1.5	0.9
No. tumors resected	–0.03	1.0	0.9–1.04	0.4
EBL (per 100 ml)	0.03	1.03	1.009–1.05	0.004
Constant	–2.8	0.07		

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