

Robot-assisted vs. Laparoscopic Partial Nephrectomy: utilization rates and perioperative outcomes

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ABSTRACT

Objectives: To examine the effect of surgical approach on perioperative morbidity and mortality after partial nephrectomy.

Materials and Methods: Within the Nationwide Inpatient Sample, patients who underwent RAPN or LPN between October 2008 and December 2009 were identified. Propensity-based matching was performed to adjust for potential baseline differences between the two groups. The rates of intraoperative and postoperative complications, blood transfusions, prolonged length of stay, and in-hospital mortality, stratified according to RAPN vs. LPN, were compared.

Results: Overall, 851 (72.5%) patients underwent RAPN and 323 (27.5%) underwent LPN. For RAPN and LPN respectively, the following rates were recorded in the propensity-score matched cohort: blood transfusions, 4.5 vs. 6.8% (p = 0.223); intraoperative complications, 5.2 vs. 2.6% (p = 0.096); postoperative complications, 10.6 vs. 13.5% (p = 0.268); prolonged length of stay, 6.8 vs. 9.4% (p = 0.238); in-hospital mortality, 0.0 vs. 0.0%. *Conclusions:* RAPN has supplanted LPN as the predominant minimally invasive surgical approach for renal masses. Perioperative outcomes after RAPN and LPN are comparable. Interpretation of these findings needs to take into account the lack of adjustment for case complexity and surgical expertise.

ARTICLE INFO

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INTRODUCTION

The incidence of renal cell carcinoma (RCC) has steadily risen in the United States in recent years. While the rates of metastatic and regional disease have remained steady, the incidence of organ confined disease has risen (1). Increased utilization of cross-sectional imaging has led to more frequent discovery of suspicious renal masses for

which surgical resection remains the standard of care (2).

Over the past decade, the management of suspicious renal masses in patients with a healthy contra-lateral renal unit has undergone a fundamental evolution; techniques developed for imperative partial nephrectomy have now become the standard of care. Due to the downward stage migration in contemporary years (1,3), utilization rate of partial nephrectomy (PN) has increased significantly (4). PN has been demonstrated to provide equivalent oncologic control (5), improved renal function, a lower risk of cardiovascular disease, as well as improved overall survival (6-8). In consequence, PN is supported by both American and European guidelines when indicated and technically feasible.

Laparoscopic PN (LPN) is a minimally invasive alternative to open partial nephrectomy (9,10). However, LPN is technically challenging, limiting its widespread adoption. Robot-assisted PN (RAPN) has emerged as a viable alternative to mitigate the technical challenges of LPN, demonstrating perioperative outcomes at least comparable to LPN, with the benefit of reduced warm ischemia time (11-13).

In the current manuscript, we explore the effect of minimally invasive surgical approach (RAPN vs. LPN) on five immediate and short-term PN outcomes. Specifically, we focus on blood transfusions, on intraoperative and postoperative complications, on prolonged length of stay (LOS) beyond the 75th percentile, and on in-hospital mortality. We rely on propensity-score matched analyses to adjust for potential differences between patients undergoing RAPN or LPN, using a contemporary (2008-2009) population-based cohort of individuals.

MATERIALS AND METHODS

Data Source

Data from the Nationwide Inpatient Sample (NIS) were abstracted between October 2008 and December 2009. The NIS includes inpatient discharge data collected via federal--state partnerships, as part of the Agency for Healthcare Research and Quality's Healthcare Cost and Utilization Project. As of the year 2009, the NIS contains data from approximately 8 million hospital stays drawn from more than 1,000 community hospitals in 44 States, approximating 20% of community hospitals within the United States, including public hospitals and academic medical centers. The NIS is the sole hospital database in the United States with charge information on all patients regardless of payer, including persons covered by Medicare, Medicaid, private insurance, and the uninsured.

Sample population and surgical procedures

Using a previously described extraction methodology (14), patients with a primary diagnosis of cancer of the kidney were identified using the International Classification of Disease, 9th Revision, Clinical Modification (ICD-9-CM) diagnostic code: 189.0; patients who underwent PN (55.4) were abstracted. Secondary diagnostic codes (ICD-9-CM 197.0, 197.7, 198.x) were used to identify patients with metastases, who were subsequently excluded from analysis. Beginning October 1, 2008, the robot-assisted modifter (ICD-9-CM 17.4x) code was recognized by the National Center of Health Statistics and the Centers for Medicare and Medicaid Services and was used to identify RAPN. Laparoscopic surgical approach was identified via concurrent procedure codes for laparoscopic exploration (54.21). The remaining patients, with an absence of RAPN or LPN codes, were determined as having undergone an OPN, and were excluded from further analysis (n = 2921).

Baseline patient and hospital characteristics

Patient characteristics include age at surgery (in years), coded as a continuous variable. Gender and race (white, black and other races including: Hispanic, Asian, Pacific Islander, Native American, or other/unknown) were also examined. Baseline Charlson Comorbidity Index (CCI) was calculated according to Charlson et al. (15), and adapted according to Deyo et al. (16). To ensure uniformity of coding across data sources, detailed insurance categories are combined in the more general groups, namely: private insurance, Medicare, Medicaid, and other (self-pay).

Hospital characteristics include location (rural vs. urban), region (Northeast, Midwest, South, West), and academic status (academic vs. non-academic), and were obtained from the American Hospital Association Annual Survey of Hospitals. Having an American Medical Association (AMA) -approved residency program, membership in the Council of Teaching Hospitals or having a ratio of full-time equivalent interns and residents to beds of 0.25 or higher defined the hospital academic status. Hospital volume was defined according to the number of partial nephrectomies performed annually.

Intraoperative complications, postoperative complications, and blood transfusions during hospitalization

The NIS records up to 15 diagnoses and procedures per in-hospital stay. The presence of any complication was defined using ICD-9 diagnoses 2 through 15. The specific ICD-9 codes used for complications relied on previously published methodology (17). Intraoperative complication was defined as accidental puncture or laceration during a procedure. Additionally, seven groups of postoperative complications were identified, namely: cardiac, respiratory, neurologic, urinary, vascular, infection, hemorrhage, operative wound, as well as septicemia (18). Blood transfusion recipients were identified using the ICD-9 procedure for transfusion of homologous blood (99.04). For purposes of statistical analysis of overall complication and transfusion rates, we stratified patients by 0 vs. 1 or greater, during hospitalization.

LOS and in-hospital mortality

LOS, provided by the NIS, is calculated by subtracting the admission date from the discharge date. In-hospital mortality information is coded from disposition of patient. Patients with missing or invalid LOS or in-hospital mortality status were not considered within the current study (n = 10). Prolonged LOS was defined as a hospitalization beyond the 75th percentile cut-off of five days.

Statistical analysis

Descriptive statistics focused on frequencies and proportions for categorical variables. Means, medians and ranges were reported for continuously coded variables. Chi-square and independent-sample t tests were used to compare the statistical significance of differences in respectively proportions and means.

Due to inherent differences between patients undergoing robotic vs. laparoscopic nephrectomy in terms of baseline patient and hospital characteristics, we relied on propensity-score matched analysis (19,20). Cohorts were matched by patient characteristics: age, gender, race, CCI and insurance status; as well as hospital characteristics: volume, location, region, and academic status. Covariate balance between the matched groups was subsequently examined. Chi-square tests were used to compare the statistical significance of differences between outcomes of RAPN and LPN.

Categorical distributions are reported as counts (%) and continuous variables as means and standard deviations (SD), and medians and interquartile ranges (IQR). Fisher's exact test, Chi-square test, and Mantel-Haenszel test were used to assess differences in distribution among categorical variables. The median test and t-test were used to assess the difference in distributions among continuous variables. All tests were two-sided with a statistical significance set a p < 0.05. Analyses were conducted using the R statistical package (the R foundation for Statistical Computing, version 2.14.1).

RESULTS

Description of sample population

Within the NIS, 1174 patients underwent minimally invasive PN between October 2008 and December 2009. Of those, 851 (72.5%) and 323 (27.5%) patients underwent RAPN and LPN, respectively. Characteristics of the patient populations are presented in Table-1a. Patient populations differed by race; a greater percentage of LPN patients were white (57.6% vs. 56.2%). No difference was noted between groups when stratification was performed according to gender, comorbidity, insurance status or income quartile. Hospital characteristics were significantly different between the two groups. Specifically, a higher proportion of RAPNs was performed at non--teaching institutions (26.2 vs. 17.6%, p = 0.002). RAPNs were more often performed in the Midwest (39.1%), whereas LPNs were most commonly performed in the Northeast (37.2%).

Propensity-score matching resulted in a cohort of 310 RAPN and 310 LPN patients (Table--1b). The standardized differences between the two groups in patient and hospital characteristics were less than 10% (19), indicating a high degree of similarity in the distribution of both

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Hospital region < 0.001
Northeast 258 (22.0) 120 (37.2) 138 (16.2)
Midwest 429 (36.5) 96 (29.7) 333 (39.1)
South 278 (23.7) 66 (20.4) 212 (24.9)
West 209 (17.8) 41 (12.7) 168 (19.7)
Hospital teaching status 0.002
Non-teaching 276 (23.8) 55 (17.6) 221 (26.2)
Teaching 882 (76.2) 258 (82.4) 624 (73.8)
Hospital volume 0.085
Mean (SD) 72 (11) 81 (121) 69 (101)
Median (IQR) 30 (13.59) 27 (12.48) 32 (14. 59)
Insurance status 0.498
Private 714 (60.8) 204 (63.2) 510 (59.9)
Medicaid 56 (4.8) 16 (5.0) 40 (4.7)
Medicare 350 (29.8) 86 (26.6) 264 (31.0)
Other 54 (4.6) 17 (5.3) 37 (4.3)

Table 1a - Demographic characteristics of patients undergoing minimally invasive partial nephrectomy – entire cohort [p values all Chi-Square].

	Overall	Laparoscopic	Robotic	р
Number of patieants	620	310	310	
Year of surgery				0.299
2008	194 (31.3)	103 (33.2)	91 (29.4)	
2009	426 (68.7)	207 (66.8)	219 (70.6)	
Gender	(),	(),	()	0.519
Male	334 (53.9)	163 (52.6)	171 (55.2)	
Female	286 (46.1)	147 (47.4)	139 (44.8)	
Age				0.633
Mean(SD)	56.3 (14.1)	56.5 (14.9)	56.0 (13.3)	
Median(IQR)	57 (48.66)	57 (47.67)	57 (48.65)	
Race				0.043
White	358 (57.7)	176 (56.8)	182 (58.7)	
Black	37 (6.0)	25 (8.1)	12 (3.9)	
Other	56 (9.0)	33 (10.6)	23 (7.4)	
Unknown	169 (27.3)	76 (24.5)	93 (30.0)	
CCI				0.983
0	419 (67.6)	208 (67.1)	211 (68.1)	
1	147 (23.7)	75 (24.2)	72 (23.2)	
2	35 (5.6)	17 (5.5)	18 (5.8)	
\geq 3	19 (3.1)	10 (3.2)	9 (2.9)	
Income				0.211
1st quartile	117 (19.4)	60 (19.9)	57 (18.9)	
2nd quartile	150 (26.3)	74 (24 5)	85 (28 1)	
3rd quartile	159 (20.3)	73 (24.3)	86 (28 5)	
4th quartile	169 (28.0)	95 (31.5)	74 (24.5)	
Hospital location			()	0 664
Rural	22 (3.5)	10 (3.2)	12 (3.9)	0.001
Urban	598 (96.5)	300 (96.8)	298 (96.1)	
Hospital region		()	()	0.002
Northeast	223 (36.0)	120 (38 7)	103 (33 2)	
Midwest	214 (34.5)	86 (27.7)	128 (41.3)	
South	108 (17.4)	66 (21.3)	42 (13.5)	
West	75 (12 1)	38 (12 3)	37 (11 9)	
Hospital teaching status	10 (12.1)	00 (12.0)	01 (11.0)	0.587
Non teaching	101 (16.2)	EQ (17 1)	10 (15 E)	
Teaching	510 (10.3)	257 (82.9)	40 (15.5) 262 (84 5)	
	515 (00.7)	207 (02.3)	202 (04.3)	0 769
Maan (CD)	94 (100)	00 (104)	00 (110)	0.700
Mean (SD)	84 (120)	83 (124)	86 (116)	
Median (IQR)	31 (13.59)	27 (11.48)	36 (16.60)	
Insurance status				0.769
Private	391 (63.1)	197 (63.5)	194 (62.6)	
Medicaid	27 (4.4)	15 (4.8)	12 (3.9)	
Medicare	173 (27.9)	82 (26.5)	91 (29.4)	
Other	29 (4.7)	16 (5.2)	13 (4.2)	

Table 1b - Demographic characteristics of patients undergoing minimally invasive partial nephrectomy – propensity-score matched cohort [p values all Chi-Square].

populations. The matched populations represent the focus of all subsequent analyses.

Intraoperative and postoperative outcomes during hospitalization

Table-2 shows the rate of intraoperative and postoperative outcomes, stratified according to RAPN and LPN, pre (Table-2a) and post-propensity score matching (Table-2b). In the post propensity--score matched cohort, the rates of intraoperative and postoperative complications were found to be equivalent, as was the rate of blood transfusion and prolonged LOS (p > 0.05). Statistically significant differences were however seen for individual complications. Specifically, patients undergoing RAPN had fewer neurologic (0.0 vs. 2.3%, p = 0.008), urinary (0.0 vs. 1.9%, p = 0.014) and bleeding complications (0.6 vs. 2.9%).

DISCUSSION

RAPN was initially described in 2004 (21) and represents an acceptable alternative to open or laparoscopic PN for selected renal masses. It has been hypothesized that RAPN may help overcome the technical challenges of LPN and may offer an easier transition to minimally invasive PN (22). In this retrospective case-control study, we compare short-term perioperative outcomes of RAPN vs. LPN, relying on a contemporary (2008-2009) population-based cohort of individuals undergoing minimally invasive partial nephrectomy. To account for baseline patient and hospital differences between the two procedure cohorts, we relied on propensity-score matched analysis to eliminate the plausible biases associated with treatment selection.

Our findings are several-fold. First, our contemporary analysis shows that RAPN has overtaken LPN as the preferred surgical approach of minimally invasive PN in the United States. In this 20% representative sample of all inpatient admissions between October 2008 and December 2009, nearly three times as many RAPNs were performed compared to LPNs (851 vs. 323), thus accounting for 72.5% of minimally invasive PNs. This finding is both novel and unexpected; the unique identifier for robotic procedures was only introduced by CMS in 2008; 2009 represents its first full year of utilization. Although objective evidence has shown that the utilization of robot--assisted radical prostatectomy has increased dramatically in recent years (23,24), no previous reported has alluded to a similar phenomenon in the context of PN.

Second, the characteristics of RAPN vs. LPN cases differed regarding several patient and hospital attributes. When compared to LPN patients, patients undergoing RAPN were less likely to be white and less likely to be treated at teaching, high-volume institutions. Moreover, LPN is predominantly performed in the Northeast, while RAPN is chiefly performed in the Midwest. LPN regionalization may be explained by the location of teaching vs. non-teaching hospital within the current database. Specifically, most teaching hospitals were located within the Northeast region, and since the majority of LPN was performed at teaching hospitals, the observed regional variability was likely a reflection of teaching vs. non-teaching hospital status. In consequence, it may be possible that lack of surgical LPN expertise in some areas could have prevented LPN being performed. In this regard, we corroborate previous findings that the use of PN is highly regionalized (25). Taken together, these findings corroborate the hypothesis that the diffusion of laparoscopy has not been uniform and is strongly influenced by patient provider characteristics, such as hospital volume and teaching status (26). Conversely, our findings also indicate that RAPN might have a different dissemination pattern, as this procedure has already been adopted in a higher proportion by surgeons at non-teaching and lower volume institutions than conventional laparoscopy. It is reasonable to presume that introduction of RAPN may in fact accelerate the utilization of PN as patients and surgeons seek to utilize minimally invasive approaches to PN. Unfortunately, temporal trend analyses are not possible since the robot-assisted modifier was only introduced in the fall of 2008.

Nonetheless, these findings emphasize the need for meticulous adjustment of patient differences when comparing perioperative outcomes. Lack of adjustment may result in important confounding of results. Specifically, based on previous

	Overall	Laparoscopic	Robotic	р
Number of patients	1174	323	851	-
Intraoperative complication	43 (3.7)	8 (2.5)	35 (4.1)	0.183
Postoperative complication				
Overall	144 (12.3)	42 (13.0)	102 (12.0)	0.635
Cardiac	14 (1.2)	3 (0.9)	11 (1.3)	0.608
Respiratory	46 (3.9)	18 (5.6)	28 (3.3)	0.072
Neurologic	17 (1.4)	7 (2.2)	10 (1.2)	0.204
Urinary	14 (1.2)	6 (1.9)	8 (0.9)	0.196
Vascular	1 (0.1)	0 (0.0)	1 (0.1)	0.538
Infection	1 (0.1)	0 (0.0)	1 (0.1)	0.538
Hemorrhage	22 (1.9)	9 (2.8)	13 (1.5)	0.155
Operative wound	8 (0.7)	1 (0.3)	7 (0.8)	0.340
Septicemia	1 (0.1)	0 (0.0)	1 (0.1)	0.538
In-hospital mortality	0 (0.0)	0 (0.0)	0 (0.0)	-
Blood transfusion	74 (6.3)	22 (6.8)	52 (6.1)	0.659
Length of stay				
Length of stay > 5 days (median)	82 (7.0)	30 (9.3)	52 (6.1)	0.056

Table 2a - Outcomes of patients undergoing minimally invasive partial nephrectomy – entire cohort (unmatched).

reports, better surgical outcomes are expected in patients treated at teaching (27) and high-volume (28) centers. To minimize bias and control for the baseline differences between RAPN and LPN patients, we relied on propensity-score matching. Propensity-score matching resulted in highly comparable patient and institutional characteristics within the RAPN and LPN groups. In propensity-score matched cohorts, no statistically significant difference between the two groups was recorded with regard to intraoperative and postoperative complications, prolonged LOS and in-hospital mortality. It is noteworthy hemorrhagic complications were significantly lower in the RAPN group (p = 0.033); though no statistically significant difference in transfusion rates was recorded (p = 0.223).

Based on the diffusion pattern of RAPN, it can be hypothesized that the utilization rates of nephron-sparing surgery will improve. Dulabon et al. (4) used a population-based dataset and identified 18330 localized RCC patients between years 1999 to 2006. PN utilization rates were 21% in 1999 vs. 45% in 2006. These findings strongly contrasts with those of Thompson et al. (29), who showed that PN rates at the Memorial Sloan-Kettering Cancer Center increased from 69 to 89% between 2000 and 2007. The disparities in reported rates highlight the difficult diffusion of LPN for small renal masses, as this technique is relatively challenging and often limited to centers of excellence.

From a practical perspective, our results indicate that on average, similar intraoperative and postoperative outcomes, including transfusion rates, prolonged LOS and in-hospital mortality, are expected whether the patient undergoes RAPN or LPN. However, these results should be interpreted with caution, since the NIS is unable to account for disease characteristics. Specifically, it is not known if complex cases, such as endophytic or hilar tumors, higher stage and grade lesions, are more often performed by the robotic or laparoscopic approach.

	Overall	Laparoscopic	Robotic	р
Number of patients	620	310	310	-
Intraoperative complication	24 (3.9)	8 (2.6)	15 (5.2)	0.096
Postoperative complication				
Overall	75 (12.1)	42 (13.5)	33 (10.6)	0.268
Cardiac	10 (1.6)	3 (1.0)	7 (2.3)	0.202
Respiratory	29 (4.7)	18 (5.8)	11 (3.5)	0.183
Neurologic	7 (1.1)	7 (2.3)	0 (0.0)	0.008
Urinary	6 (1.0)	6 (1.9)	0 (0.0)	0.014
Vascular	0 (0.0)	0 (0.0)	0 (0.0)	-
Infection	0 (0.0)	0 (0.0)	0 (0.0)	-
Hemorrhage	11 (1.8)	9 (2.9)	2 (0.6)	0.033
Operative wound	1 (0.2)	1 (0.3)	0 (0.0)	0.317
Septicemia	1 (0.2)	0 (0.0)	1 (0.3)	0.317
In-hospital mortality	0 (0.0)	0 (0.0)	0 (0.0)	-
Blood transfusion	35 (5.6)	21 (6.8)	14 (4.5)	0.223
Length of stay				
Length of stay > 5 days (median)	50 (8.1)	29 (9.4)	21 (6.8)	0.238

Table 2b - Outcomes of propensity-score matched patients undergoing minimally invasive partial nephrectomy. Patients were matched by age, gender, race, CCI and insurance status; as well as by hospital characteristics: volume, location, region, and academic status.

To the best of our knowledge, the current manuscript represents the first population-based report to compare complication rates after RAPN and LPN. Benway et al. recently published the largest multi-institutional comparison of the two approaches, describing the outcomes of three experienced minimally invasive surgeons (n = 247) (11). They found similar post-operative complication rates of RAPN vs. LPN (8.6 vs. 10.2%), which corroborate the findings of this study.

A significant limitation of the NIS is the lack of capture of tumor stage and pathological characteristics; there was likely some variability in distribution and this would affect our findings. There may have also been unobserved differences in operative characteristics that were not captured in discharge level data, ie. warm ischemia time. Although propensity-based matching was performed, there may have also been differences amongst the groups concerning important patient variables (personal preferences, education, body mass index and medication utilization), as well as socio-economical determinants. Moreover, our mortality estimates are based on in-hospital rates. It is possible that the true mortality is underestimated as some patients may have died at other institutions where their mortality was not captured. Finally, we were not able to classify complications according to ideal, prospective methodologies, such as the Clavien classification, nor were we able to grade them using the Common toxicity Criteria for Adverse Events (30,31).

CONCLUSIONS

RAPN has supplanted LPN as the predominant minimally invasive surgical approach for renal masses. Perioperative outcomes after RAPN and LPN are comparable and remain comparable following adjustment for several common confounders. Interpretation of these findings needs to take into account the lack of adjustment for case complexity and surgical expertise.

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CONFLICT OF INTEREST

None declared.

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