

Robot-assisted adrenalectomy: current perspectives

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Abstract: Laparoscopy has established itself as the procedure of choice for performing adrenalectomy in benign adrenal disorders. Although laparoscopy scores heavily over open approach in terms of lesser blood loss, pain, shorter hospital stay and better cosmesis, it is riddled with certain shortcomings such as the need of dexterity, two-dimensional vision, dependence on an assistant for camera, etc. Robotic surgery promises to overcome these limitations. Multiple series have established that robotic adrenalectomy is a safe and effective procedure as conventional laparoscopy. Recently, robotic surgery has been found to be precise and accurate in performing partial adrenalectomy in hereditary adrenal syndrome cases. Other advances like single-port surgery have expanded the horizon and indications of robotic surgery. This review aims at studying the current evidence available for the effectiveness of robot-assisted adrenalectomy and defining its current status in managing adrenal disorders.

Keywords: robotics, adrenal gland, adrenalectomy, laparoscopy, pheochromocytoma

Introduction

Laparoscopic adrenalectomy (LA) has been recognized as the procedure of choice for the management of, especially benign, adrenal disorders. Various studies have proved the feasibility, efficacy and safety of this approach.¹⁻⁴ As compared to open adrenalectomy (OA), LA involves lesser blood loss, lesser pain and shorter hospital stay and offers better cosmesis.⁵⁻¹² Despite all these advantages, LA has been restricted to a handful of experienced surgeons at high-volume centers because of certain technical drawbacks like two-dimensional view, stiff non-articulating instruments, dependence on an assistant for camera, etc. Robot-assisted adrenalectomy (RA) is the recent addition to the armamentarium of surgeons. RA overcomes the limitations of laparoscopic surgery by providing a three-dimensional magnified view, better ergonomics, control of the camera and multi-articulated instruments using Endowrist technology to the surgeons. Recent studies, reviews and meta-analyses have tried to establish its superiority over LA and OA in terms of faster convalescence and lesser complication rate.¹³⁻²⁵ Higher cost and longer operating times have although prevented RA from gaining widespread acceptance. A recent large international series has shown that conventional laparoscopy and laparoendoscopic single-site surgery seem to be the most common adopted techniques, whereas minilaparoscopy and RA seem to be gaining popularity at a slower rate.²⁶ In this review, the current status of robotic approach in performing adrenalectomy has been analyzed.

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Feasibility

In 2000, Horgan and Vanuno first reported the use of robot in performing laparoscopic bilateral adrenalectomy.²⁷ Since then, many authors published their experience of performing robot-assisted LA in small series of patients until Winter et al published the first series on 30 patients in 2006. Three different surgeons at a single center performed the surgeries. The median operative (OR) time was 185 minutes, postoperative complication rate was 7% and median hospital stay was 2 days and there was no conversion to open or laparoscopic surgery.²⁸ Since then, more than 50 studies have been published describing RA, but few studies have included more than 20 cases. This is the minimum number of cases required for achieving the learning curve of RA.¹⁶

Table 1 describes the various case series that have evaluated the feasibility of robotic adrenalectomy (>20 patients). Subsequent series have shown comparatively decreasing OR time with decreasing conversion rate. Multiple studies in recent past comparing robotic adrenalectomy with LA have shown nil conversion rate.

RA vs LA

Multiple series have compared laparoscopic and robotic transperitoneal adrenalectomy. Most of these studies have been retrospective observational comparative studies,^{13,14,16–24,26} and

only one of them was a randomized controlled trial.¹⁵ One of the first series comparing LA with robotic adrenalectomy was by Morino et al who evaluated outcomes in 10 patients who underwent laparoscopic and robotic adrenalectomy.¹⁵ The authors observed that although the laparoscopic approach took less time (115 vs 169 minutes), it had a higher conversion rate (4 vs 0) and longer length of stay (LOS; 5.7 vs 5.4 days) as compared to robotic approach. Brunaud et al performed a retrospective analysis of a large series of 50 robotic and 59 laparoscopic adrenalectomies. They found that though the OR time was longer in the robotic group (189 vs 159 minutes), it was associated with a lesser blood loss (49 vs 71 mL) and shorter LOS (6.3 vs 6.9 days) with similar conversion rate (4 vs 4).¹⁶ Table 2 describes various studies that have compared the outcomes in laparoscopic and robotic groups.

Brandao et al recently conducted a systematic review and meta-analysis comparing laparoscopic and robotic adrenalectomy.²⁰ This meta-analysis of nine studies, eight retrospective observational and one randomized controlled trial, included 600 patients (277 RA vs 323 LA). Although they did not observe any difference in terms of conversion rates, OR time, or postoperative complications, the robotic group was found to have less blood loss and a short hospital stay. As a part of the International Consultation on Urologic Diseases and European Association of Urology consultation on Minimally Invasive Surgery in Urology, an extensive methodological systemic review of the literature of laparoscopic and robotic adrenalectomy in the treatment of adrenal diseases was performed. The literature was searched systematically till January 2014 to identify studies comparing the safety and efficacy of different modalities of minimally invasive adrenal surgery techniques. The authors presented major findings in an evidence-based fashion and

Table 1 Robotic adrenalectomy feasibility series

Study	No. of patients	OR time (minutes)	Conversion rate (%)	Morbidity (%)
Brunaud et al ²⁹	100	95	5	8
Giulianotti et al ³⁰	41	118	0	4.8
Nordenström et al ³¹	100	113	7	13
Raman et al ³²	40	117	4	10
Pahwa et al ³³	25	139 ± 30	0	12

Abbreviation: OR, operative.

Table 2 Studies comparing LA and RA

Study	No. of patients (RA vs LA)	OR time (minutes) (RA vs LA)	EBL (mL) (RA vs LA)	Conversion (RA vs LA)	Complications (RA vs LA)
Morino et al ¹⁵	10 vs 10	169 vs 114	NR	4 vs 0	0 vs 0
Brunaud et al ¹⁶	50 vs 59	189 vs 159	49 vs 71	4 vs 4	5 vs 9
Agcaoglu et al ¹⁴	24 vs 38	159 vs 187	84 vs 167	1 vs 4	0 vs 0
Agcaoglu et al ¹⁷	31 vs 31	163 vs 166	25 vs 36	NR	0 vs 0
Karabulut et al ¹³	50 vs 50	166 vs 164	41 vs 41	1 vs 2	1 vs 5
Pineda-Solis et al ²¹	30 vs 30	190 vs 160	30 vs 55	0 vs 5	0 vs 0
Aksoy et al ¹⁸	42 vs 57	186 vs 187	50 vs 77	0 vs 3	1 vs 2
You et al ²²	15 vs 8	207 vs 183	NR	0 vs 0	2 vs 2
Aliyev et al ¹⁹	25 vs 40	149 vs 178	36 vs 43	1 vs 3	1 vs 2
Brandao et al ²³	30 vs 46	120 vs 120	50 vs 100	0 vs 1	7 vs 11
Pavan et al ²⁶	80 vs 337	150 vs 120	50 vs 50	2 vs 10	22 vs 40
Lairmore et al ²⁴	66 vs 17	152.8 vs 177.3	103.4 vs 46.5	8 vs 1	15 vs 11.8

Abbreviations: LA, laparoscopic adrenalectomy; RA, robot-assisted adrenalectomy; OR, operative; EBL, estimated blood loss; NR, not recorded.

provided a set of recommendations. They concluded that RA might be considered an alternative to LA but requires further study (Grade B).³⁴

Tang et al performed a meta-analysis that included eight studies (232 cases and 297 controls) assessing RA vs LA, where six were prospective and two were retrospective. Patients in the LA group had significantly shorter OR time (weighted mean difference [WMD] = 17.52 minutes; 95% confidence interval [CI], 3.48–31.56; $p = 0.01$), but patients in the RA group had significantly lesser estimated blood loss (EBL; WMD = –19.00 mL; 95% CI, –34.58 to –3.41; $p = 0.02$) and shorter LOS (WMD = –0.35 day; 95% CI, –0.51 to –0.19; $p < 0.001$). Patients in both the groups had similar conversion rates and overall complications.³⁵ The same findings have been corroborated in another meta-analysis and systematic review by Chai et al.³⁶

RA has been compared with conventional LA in a certain subset of patients too. Aksoy et al compared outcomes of these two approaches in obese patients (42 patients in robotic group and 57 in laparoscopic group). The authors observed similar OR times (186.1 vs 187.3 minutes), less EBL (50.3 vs 76.6 mL) and shorter LOS (1.2 vs 1.7 days) in the robotic group. There were no conversions in the robotic group, whereas the conversion rate was 5.2% in the laparoscopic group.¹⁸ Aliyev et al compared perioperative outcomes and efficacy of laparoscopic and robotic adrenalectomies in patients with pheochromocytoma. Patients in robotic group had a shorter OR time (149 vs 178 minutes), less EBL (36 vs 43 mL), fewer conversions (1 vs 3) and shorter LOS (1.2 vs 1.7 days).¹⁹ Agcaoglu et al performed a prospective comparative study of 24 patients in robotic group with 38 patients in laparoscopic group undergoing adrenalectomy for large tumors (mean size: 6.5 ± 0.4 [robotic] vs 6.2 ± 0.3 cm [laparoscopic]). The robotic group had a shorter OR (159.4 ± 13.4 vs 187.2 ± 8.3 minutes, $p = 0.043$), shorter LOS (1.4 ± 0.2 vs 1.9 ± 0.1 days, $p = 0.009$) and lesser conversion rate (4% vs 11%, $p = 0.43$) but similar EBL ($p = 0.147$). The perioperative morbidity was nil in the robotic group and 2.7% in the laparoscopic group.¹⁴

Studies comparing RA with OA are less. Probst et al recently published their trial comparing RA with OA. They analyzed 28 matched pairs of patients with respect to their characteristics, perioperative outcomes and cost effectiveness. Patients in RA group benefited from a shorter LOS (11.1 ± 4.8 vs 6.8 ± 1.2 days, $p < 0.01$) and intermediate care treatment (2.3 ± 1.7 vs 1.2 ± 0.4 days, $p < 0.01$). The mean operating time was longer for RA (128.5 ± 46.5 vs 102.2 ± 44.5 minutes, $p = 0.03$), but the OR time of the last 10 RA

procedures (mean: 97.1 ± 35.2 minutes) was similar to that of OA. There was no difference in the rate of complications between the two groups.³⁷

Posterior retroperitoneoscopic adrenalectomy (PRA)

Theoretically, posterior retroperitoneal approach of adrenalectomy offers distinct advantages like decreased chance of postoperative ileus and other intestinal complications, as peritoneum is not breached, along with decreased postoperative pain. Ludwig et al first reported on their experience of performing robotic PRA on six patients. The mean tumor size in the study population was 2.8 cm with a mean OR time of 121 minutes. The OR time was further reduced to a mean of 57 minutes for five patients where the entire dissection was performed robotically. The authors did not observe any morbidity or mortality.³⁸ Berber et al published their experience of performing RA on 23 patients, of whom eight underwent retroperitoneal adrenalectomy. The average tumor size in this study was 2.9 cm with an average OR time of over 3.5 hours. More importantly, the docking time decreased to 15 minutes in the last four cases from 1 hour in the beginning of the study. According to the authors, patients with smaller tumors, bilateral disease and who have undergone multiple prior operations are best suited for retroperitoneal approach.³⁹ The surgeons in both the series had a vast experience of performing retroperitoneoscopic surgery and robotic procedures.

Karabulut et al compared robotic adrenalectomy with the laparoscopic approach. Thirty-two patients were operated through transabdominal route and 18 were operated through posterior retroperitoneoscopic approach in both the laparoscopic and robotic groups. They did a step-by-step evaluation of OR time and found that both the LA and robotic PRA had similar OR times for each step except for shorter time for hemostasis in the robotic group (23 ± 4 vs 42 ± 9 minutes, $p = 0.03$). Later on, the same group published their results when they did a head-to-head comparison of laparoscopic and robotic PRA on 31 patients each. Although the mean (standard error of the mean, SEM) OR time was similar in both the groups, there was a significant difference in the mean (SEM) OR time of the last 21 cases of robotic-assisted PRA when compared to the laparoscopic group ($139.1 [10.9]$ vs $167.7 [12.1]$ minutes, respectively; $p = 0.046$). The mean (SEM) EBL was similar in the robotic and laparoscopic groups ($25.3 [10.2]$ and $35.6 [9.9]$ mL, respectively; $p = 0.24$). The median hospital stay was 1 day for both groups. Both the groups had similar overall postoperative narcotic use.¹³

Lairmore et al performed a retrospective analysis of their minimally invasive adrenalectomy patients operated over the last 10 years by three different approaches. The study included 67 transperitoneal LA (TLA), 76 PRA and 17 robot-assisted PRA (RAPRA) procedures. Tumor size in patients undergoing retroperitoneal adrenalectomy was smaller than in patients undergoing TLA (2.38 vs 3.6 cm, $p < 0.0001$). Procedure time was significantly longer in the robotic group as compared to TLA/PRA (177 vs 153/133 minutes, $p = 0.008$), but LOS significantly decreased (1.53 vs 2.82/1.85 days, $p = 0.004$). Mean EBL was also lower for RAPRA than for LA (46 ± 25.4 vs 103.4 ± 153.8 mL, $p = 0.005$).²⁴

RA for malignant diseases

The use of minimally invasive approach for adrenocortical carcinoma is still a matter of debate. There has not been any prospective or randomized controlled trial comparing robotic or laparoscopy technique with open surgery. Some retrospective case reports and series have reported increased recurrence, peritoneal carcinomatosis, positive margins and local recurrence rates for laparoscopic cases compared to open surgery.^{40–44} Robotic adrenalectomy has so far been mainly performed for benign diseases, but it has also been reported for adrenal cancer, adrenal metastasis and oncocytoma.^{45–47}

Recent advances in robotic adrenalectomy

Single-port surgery

Park et al first described the use of robot for performing single-port PRA. Five patients underwent the procedure. The mean tumor size, OR time and EBL were 1.48 ± 0.28 (range: 1.0–1.7) cm, 159.4 ± 57.6 (range: 103–245) minutes and 46.0 ± 56.8 (range: 5–120) mL, respectively. The average time to oral intake and postoperative hospital stay was 0.65 ± 0.11 (range: 0.54–0.79) days and 4.0 ± 2.23 (range: 3–8) days, respectively. There were no conversions to open surgery or postoperative complications.⁴⁸ Arghami et al described their experience of performing single-port RA in 16 patients and did a matched cohort analysis with 16 patients of LA. The OR time was 183 ± 33 minutes for single-port RA and 173 ± 40 minutes for LA ($p = 0.58$). There was one conversion to OA (6%) in each group, both because of bleeding on the right side during bilateral adrenalectomy. Two right-sided single-port RA patients required conversion to LA, one because of poor visualization. Both groups had similar pain scores (mean of 3.7 on a scale from 1 to 10) on postoperative day (POD) 1, and patients in the single-port RA used less

narcotic pain medication in the first 24 hours after surgery (43 vs 84 mg in LA group, $p = 0.001$). The differences between the single-port RA group and LA group in LOS (2.3 ± 0.5 vs 3.1 ± 0.9 days, $p = 0.23$), percentage of patients discharged on POD 1 (56% vs 31%, $p = 0.10$) and hospital cost (16% lower in single-port RA group, $p = 0.17$) did not reach statistical significance.⁴⁹

Lee et al described their experience of performing robotic single-site adrenalectomy in 33 patients. The mean OR time was 118 ± 25.8 minutes. Sixty-seven percent of patients had pain scores of less than 4 (on a scale of 1–10). Seventy-four percent of patients were discharged on POD 1, and 96% were discharged on POD 2. OR times were found to drop significantly from a mean of 124 to 103 minutes after 21 adrenalectomies ($p = 0.05$).⁵⁰

Robotic partial adrenalectomy

Although total adrenalectomy has been traditionally advocated for bilateral adrenal disorders especially in hereditary syndromes like multiple endocrine neoplasia type 2, Von Hippel–Lindau disease and neurofibromatosis type I with decreased chances of recurrence, its benefits must be weighed against the morbidity of medical adrenal replacement therapy. Lifelong adrenal replacement therapy after bilateral adrenalectomy may predispose patients to osteoporosis, Addisonian crisis and decreased quality of life.⁵¹ Hence, adrenal-sparing surgery or partial adrenalectomy has been suggested for patients with hereditary adrenal-producing syndromes, bilateral or multifocal lesions or solitary adrenal glands.

Kumar et al first described the technique of robotic-assisted partial adrenalectomy in a patient with isolated adrenal metastasis.⁴⁷ Boris et al described their initial experience of 13 partial adrenalectomies in 10 patients. Median OR time was 200 minutes, median blood loss was 150 mL and median tumor size was 2.7 cm. No patient developed any intraoperative complication related to catecholamine surge like hypertensive crisis, prolonged hypotension, myocardial infarction or cerebrovascular accident. There were no recurrences, and only one patient required steroid replacement.⁵²

There have been no comparative or prospective series of robotic and laparoscopic partial adrenalectomy. Hence, more research is required to fully define the role of robotic-assisted partial adrenalectomy.

Cost

Cost of robotic surgery is one of the prime and significant concerns for its widespread acceptance globally. The high

cost associated with robot is mainly related to its high procurement cost, use of expensive surgical consumables and longer OR times along with high maintenance cost. Hence, a significant advantage needs to be established over other approaches so as to overcome this cost barrier. In general, cost of surgical consumables along with maintenance charges per procedure roughly adds 900–950\$ to the usual cost of the surgery.¹⁴ Another study estimated the additional cost to be around 1400–2900\$ for performing unilateral RA as compared to LA.⁵³ A study by Bodner calculated that RA was approximately 1.5 times more costly than LA.⁵⁴ On the contrary, Winter et al reported no significant difference between RA and OA or LA in terms of total cost.²⁸ In another series, RA was found to be 2.3 times more expensive than LA. The authors suggested that this cost difference could be offset by multidisciplinary and high-volume use of robots by other surgical specialties and when depreciation of robotic equipment is distributed over 10 years instead of 5 years.²⁹ Arghami et al showed that single-port RA was 16% less expensive than LA ($84\% \pm 14\%$ vs $100\% \pm 16\%$), although the difference was not statistically significant.⁴⁹ Probst et al calculated that the additional cost of robotic procedure is €2288 per procedure provided there are more than 150 robotic procedures in a year. The expenses could be further reduced if there are more cases in high-volume centers. They also showed that the overall cost for patients undergoing RA was lower than that for patients undergoing OA because of difference in the length of hospital stay.³⁷

Conclusion

Robotic approach is safe, feasible and as effective as conventional laparoscopy for performing adrenalectomy, especially for benign adrenal disorders. Depending upon the body habitus of the patient and experience of the surgeon, either transperitoneal route or retroperitoneal approach can be safely used to perform RA. Robotic surgery has also been demonstrated to be superior to laparoscopy in cases of large tumors, partial adrenalectomy and pheochromocytomas and in obese patients, but the number of patients has been limited in these retrospective series. Higher cost and lesser number of patients preclude the widespread adaption of this technique. High-volume centers and experienced surgeons can safely adapt this technology for better ergonomics of the surgeons and for better and precise dissection of adrenal tumors. Well-designed randomized controlled trials can clearly establish the exact status of robotics in performing radical adrenalectomy.

Disclosure

The author reports no conflicts of interest in this work.

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