



Surgical approach to patients with hypercortisolism

Colleen M. Kiernan, Carmen C. Solórzano

Vanderbilt University Medical Center, Nashville, TN, USA

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Correspondence to: Colleen M. Kiernan; Carmen C. Solórzano. Vanderbilt University Medical Center, Nashville, TN, USA.

Email: colleen.m.kiernan@vumc.org; carmen.solorzano@vumc.org.

Abstract: According to the Endocrine Society Clinical Practice Guidelines, the goal of treating overt Cushing's syndrome is to control cortisol levels or more importantly its actions at its receptor in order to eliminate the end organ effects and treat comorbidities associated with hypercortisolism. This chapter will review the surgical management of hypercortisolism. It will be subdivided into two main sections: the management of: (I) ACTH-dependent; and (II) ACTH-independent (adrenal) hypercortisolism. The perioperative factors that surgeons should consider after the diagnosis has been made will also be discussed. Lastly, the utilization of robotic surgery for adrenalectomy and the perceived benefits and potential pitfalls of this approach when treating patients with hypercortisolism will be reviewed.

Keywords: Hypercortisolism; adrenal; Cushing's disease (CD); Cushing's syndrome; adrenalectomy

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ACTH-dependent hypercortisolism

In ACTH-dependent hypercortisolism, treatment focuses on the surgical removal of the primary source of ACTH production. However, when the source is occult, surgical removal is not feasible or resection and/or debulking fails to control the disease, bilateral adrenalectomy can be considered.

Pituitary ACTH-dependent hypercortisolism

Hypercortisolism caused by an ACTH-secreting pituitary adenoma (Cushing's disease, CD) is the most common cause (70%) of endogenous hypercortisolism (1). The goals of treatment are to achieve remission of disease and long-term control without recurrence. Transsphenoidal microsurgery is the recommended first line treatment for pituitary overproduction of ACTH (2). Transsphenoidal surgery (TSS) achieves remission in 42–97% of cases (1). In cases in which surgery fails, repeat TSS, radiotherapy, medical therapy, and/or bilateral adrenalectomy are options for

management. Extensive discussion of pituitary reoperation and/or medical therapy of CD are beyond the scope of this chapter.

Bilateral adrenalectomy for CD has nearly a 100% success rate and results in immediate control of steroid excess. However, it results in lifelong dependency on glucocorticoid substitution and the risk of developing Nelson's syndrome and therefore should only be considered when all other management options have been exhausted for persistent or recurrent CD. Nelson's syndrome is a rare clinical manifestation that occurs in 8–47% of patients after bilateral adrenalectomy. It is associated with the clinical triad of hyperpigmentation, excessive ACTH secretion, and a pituitary adenoma (3). *Table 1* describes the management options available in patients with CD after failure of TSS as well as the perceived advantages and disadvantages of each option.

Ectopic ACTH-dependent hypercortisolism

One of the more challenging causes of hypercortisolism

Table 1 management options for Cushing's disease after failure of transsphenoidal surgery and medical management

Options	Advantages	Disadvantages
Second/Third Transsphenoidal Surgery	Persistent remission 37–70% (4)	Hypopituitarism 5–50% Adrenal insufficiency and crisis
Radiation Therapy	40–60% success rate (4)	Delayed effect Hypopituitarism 21–57% Cranial nerve palsy Adrenal insufficiency and crisis
Medical Therapy	25–90% success rate (5) Can delay the need for invasive treatment	Life-long therapy Medication side effects (nausea, vomiting, transaminitis, hirsutism) Adrenal insufficiency and crisis
Bilateral Adrenalectomy	<ul style="list-style-type: none"> • Nearly 100% success rate • Immediate control of steroid excess • Improved health related quality of life in ~80% of patients (5) 	Adrenal insufficiency and crisis Nelson's tumor Surgical morbidity 18% Surgical mortality 3% (6)

for the adrenal surgeon is ectopic ACTH-dependent hypercortisolism or ectopic ACTH Cushing's syndrome (EAS). EAS comprises 5% to 20% of all cases of ACTH-dependent hypercortisolism (7-9). EAS is commonly associated with neuroendocrine tumors (NETs) such as bronchial, pancreatic and thymic carcinoids, as well as, small cell lung carcinomas, medullary thyroid carcinoma, and pheochromocytomas. However, occult tumors occur in approximately 12–38% of the reported series making EAS difficult to treat because the primary tumor cannot be localized (7).

The primary treatment goal in patients with EAS is to remove the source of autonomous ACTH without permanent disruption of the pituitary-adrenal axis (4). When the ectopic source of ACTH is identified the optimal treatment is extirpation of the primary or metastatic tumor. In patients with a single primary tumor, resection has been associated with an 83% remission rate (7,10,11). Overall, attempts at removal of the ACTH tumor source fails in over 50% of patients, partly due to unresectable metastases and also because many tumors are small and therefore difficult to localize (12). Thus, a substantial number of patients (30–56%) ultimately undergo bilateral adrenalectomy when the EAS source remains occult and/or the hypercortisolism cannot be treated by other means (7,10,11).

Prior to proceeding with bilateral adrenalectomy for the management of EAS, every effort should be made to localize the primary or metastatic source. As these tumors can occur in multiple sites, thin-cut cross sectional imaging (CT or MRI) of the chest, abdomen and pelvis is recommended as

the initial imaging modality (7). However the sensitivity for detection for the source of EAS is low, ranging from 52–66% (11,13). Whereas previous studies utilizing [(18)F] fluorodeoxyglucose (FDG) positron emission tomography (PET) imaging have concluded that FDG-PET was not useful in localizing tumors that were occult on CT or MRI (14), others have concluded that FDG-PET was helpful in localizing the source of ACTH in small-cell lung NETs, atypical thymic tumors and medullary thyroid carcinoma but not helpful in well differentiated or low grade NETs (15).

Recently, in a multi-institutional retrospective series of 28 patients with EAS the use of PET/CT with [⁶⁸Ga] - DOTATATE (DOTA - (Tyr³) - octreotate) was shown to localize the primary tumor and/or metastases or change management in the majority of study patients (18/28, 64%). In this study, 11/17 (65%) of previously occult tumors were identified using [⁶⁸Ga] - DOTATATE (12). Thus, [⁶⁸Ga] - DOTATATE may prove to be a useful adjunct in localizing and managing previously occult ACTH producing tumors. Consideration of additional organ specific imaging such as thyroid ultrasound or endoscopic ultrasound of the pancreas can be entertained based upon clinical suspicion, the results of other imaging studies (CT, MRI, [⁶⁸Ga] - DOTATATE or FDG PET/CT) results or biochemical evaluation. To our knowledge the current guidelines (ENDO and NCCN) have not clearly recommended an imaging algorithm for AES patients.

Once all localizing modalities have been exhausted and/or medical management or surgical resection/debulking of

the primary or metastatic tumor has failed, surgical bilateral adrenalectomy may be reasonable in select patients. Ideally, this is performed in an elective setting, however occasionally in severe EAS the need for adrenalectomy becomes more urgent. This is typically encountered when a patient's cortisol elevation results in significant end organ effects such as sepsis, severe and intractable hypokalemia, uncontrolled hypertension, heart failure, gastrointestinal hemorrhage, glucocorticoid-induced acute psychosis, progressive debilitating myopathy, thromboembolism, and/or uncontrolled hyperglycemia (4). It is essential to stabilize these patients and manage the end organ effects prior to proceeding to the operating room. Medical management of hypercortisolism may include a combination of mitotane, metyrapone, ketoconazole and/or etomidate to inhibit steroidogenesis.

Bilateral adrenalectomy for ACTH-dependent hypercortisolism operative considerations

Bilateral adrenalectomy is currently considered a second or third line treatment for ACTH-dependent hypercortisolism, either CD or EAS (16). Since the introduction laparoscopic adrenalectomy in 1992, minimally invasive surgery (MIS) of the adrenal gland has become the preferred approach for bilateral adrenalectomy. Minimally invasive approaches include laparoscopic transabdominal (LTA), posterior retroperitoneoscopic (PRA), and robotic-assisted TA and PRA (RTA or RPRA). MIS has been associated with less blood loss and shorter hospital stays when compared to open approaches (17). While no individual MIS approach has been demonstrated to be superior, there are certainly advantages and disadvantages to each (Table 2). A review of 23 studies that included 739 patients who underwent bilateral adrenalectomy (approximately 70% performed laparoscopically) reported a surgical morbidity of 18% and 30-day mortality of 3% (6).

While some centers advocate for bilateral PRA to avoid patient repositioning, in our experience many of these patients are better candidates for bilateral TA adrenalectomy which can be performed laparoscopic or robotic safely and efficiently. The PRA approach can be difficult in this patient population for two reasons: (I) central obesity, abdominal wall thickness and the marked increase in retroperitoneal fat making access and visualization difficult and (II) the majority of these patients are medically unfit and may not tolerate the prone position for a prolonged period. The authors also feel that the TA approach may allow for a more complete

wide adrenalectomy including a generous amount of *en-bloc* periadrenal fat given the more familiar anatomy and less constricted space. Failure to remove all adrenal tissue in these patients leads to persistent hypercortisolism. PRA remains a feasible approach in a select group of these patients.

Bilateral adrenalectomy for ACTH-dependent hypercortisolism postoperative considerations

The resection of both adrenal glands causes permanent adrenal insufficiency resulting in the lifelong need for glucocorticoid and mineralocorticoid replacement. Patients are typically given "stress-dose" steroids in the perioperative period. In our institution we use hydrocortisone 50 mg intravenously every 8 hours. Once the patient is tolerating oral intake they are transitioned to oral hydrocortisone. Most patients with adrenal insufficiency require 15–25 mg of hydrocortisone daily in split doses. The first dose should be taken when the patient first wakes up and the last dose in the evening (19). Most patients should take 50–200 µg of fludrocortisone as a single daily dose. If essential hypertension develops, the dose of fludrocortisone can be reduced, but not stopped (19). Surgery, invasive medical procedures or physiologic stress require administration of additional stress dose steroids.

The risk of adrenal crisis must be discussed with patients and ample education must be provided regarding signs and symptoms of adrenal insufficiency and adrenal crisis and the necessity of glucocorticoid and mineralocorticoid compliance and dose adjustment in times of stress. Medical alert jewelry should be required to alert medical personnel in case of loss of consciousness occurs in a time of adrenal insufficiency. Additionally, patients should have additional doses of hydrocortisone tablets as well as an injectable hydrocortisone formulation in case of emergency. Patients should be instructed to double the daily hydrocortisone dose in times of illness/stress and notify health care providers if they are unable to tolerate PO medications due to gastrointestinal disturbances.

Adrenal crisis is a life-threatening emergency that requires immediate recognition and treatment. Approximately 8% of patients with adrenal insufficiency will experience an adrenal crisis each year (20–22). This most commonly occurs during a gastrointestinal illness but can also be precipitated by infection, invasive procedures, injuries, physical activity, pregnancy and psychological stress (23). Prompt medical attention with immediate intramuscular or intravenous high dose hydrocortisone

Table 2 Commonly utilized minimally invasive operative approaches to the adrenal gland

Operative approach	Advantages	Disadvantages	Patient factors to consider
Laparoscopic transabdominal adrenalectomy (LTA)	<ul style="list-style-type: none"> Large working space Familiar orientation of anatomy 	<ul style="list-style-type: none"> Requires mobilization of intraabdominal organs Must reposition for bilateral adrenalectomy 	<ul style="list-style-type: none"> Useful in morbidly obese patients vs. open May be difficult in patients with prior abdominal surgery Large fatty liver
Posterior retroperitoneoscopic adrenalectomy (PRA)	<ul style="list-style-type: none"> Direct access to gland Avoids intra-abdominal cavity No reposition for bilateral adrenalectomy 	<ul style="list-style-type: none"> Lack of access to abdomen for exploration or control of hemorrhage Difficulty in removing large tumors Can be difficult in obese patients Decreased working space 	<ul style="list-style-type: none"> Distance from skin to fascia and gland in obese patients Inability to tolerate prone positioning Excess retroperitoneal adiposity
Robotic transabdominal (RTA) or robotic posterior retroperitoneoscopic (RPRA)	<ul style="list-style-type: none"> Useful in Obese and super obese patients (RTA) Can be used for larger tumors (>5.5 cm) (RTA) 3D depth perception, additional dexterity and fine dissection (both) Dexterity in tighter space and allowing less tumor manipulation (RPRA) Ergonomics 	<ul style="list-style-type: none"> Perceived higher cost Complexity 	<ul style="list-style-type: none"> Similar to patient factors to consider for LA or PRA

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(100 mg) and intravenous fluids is essential.

It is also important for clinicians caring for patients after bilateral adrenalectomy to note that the preoperative hypercortisolism can impact the patient's immunity and increase the risk of postoperative infectious complications. Additionally hypercortisolism alters the coagulation cascade for up to 1 year after surgical cure (16,24) and is associated with increased risk of deep vein thrombosis (DVT),

especially in the first postoperative month (24-26). Thus prophylactic anticoagulation should be considered.

Monitoring for persistent or recurrent disease is an important aspect of postoperative management of EAS and CD patients undergoing bilateral adrenalectomy. In a review of 23 studies that included 739 patients who underwent bilateral adrenalectomy 3-34% patients had residual cortisol secretion due to accessory adrenal tissue

or adrenal remnants, but less than 2% had a relapse of hypercortisolism (6). Rarely, residual adrenal cortical tissue regrows in the surgical bed or even more rarely in the gonads as a result of prolonged ACTH hypersecretion. If a patient develops signs or symptoms of hypercortisolism after bilateral adrenalectomy, clinicians should test endogenous cortisol secretion by withholding steroid replacement for 24 hours and checking morning serum cortisol levels (16).

ACTH-independent hypercortisolism

The majority ACTH-independent or adrenal based cases of hypercortisolism are due to unilateral adrenal adenomas. Much less frequently occurring etiologies of ACTH-independent hypercortisolism are bilateral adrenal macronodular hyperplasia (BAMH), primary pigmented adrenal disease (PPAD) and adrenal cortical carcinoma (ACC). In this section we will discuss the preoperative, operative and postoperative considerations for adrenal surgeons treating ACTH-independent hypercortisolism.

Unilateral adrenal adenoma

Unilateral adrenal adenomas are benign neoplasms of the adrenal gland that account for approximately 60% of adrenal causes of hypercortisolism (27). They are often incidentally discovered on cross sectional imaging but may also be localized after a patient presents with signs and symptoms or Cushing's syndrome and biochemical evaluation is consistent with an adrenal source of hypercortisolism.

After the diagnosis has been confirmed and the gland has been localized with cross-sectional imaging, the recommended management for adrenal hypercortisolism is unilateral adrenalectomy (see below for exceptions and rare conditions).

As previously discussed, most patients with adrenal tumors can undergo a MIS. Such approach is associated with shorter hospitalization, less pain and morbidity, quicker recovery, and low incidence of incisional hernias. An absolute contraindication to MIS adrenalectomy is extension of the tumor into nearby structures. A relative contraindication is tumor size when ACC is highly suspected. From a practical standpoint, tumors >6 to 8 cm may be more difficult to mobilize and manipulate and can often result in conversion to an open procedure. Additionally, as discussed in the subsequent section adrenal tumors >4–6 cm that produce cortisol and are indeterminate on cross sectional imaging

warrant careful consideration of the operative approach due to the increased risk of ACC.

BAMH

BAMH is very rare cause of ACTH independent hypercortisolism. It presents on imaging with characteristic multiple bilateral macronodules (>10 mm) with hyperplasia and/or internodular atrophy. It is often diagnosed as bilateral adrenal incidentalomas with autonomous cortisol secretion (ACS) (28,29).

The treatment of choice to control hypercortisolism in patients with BAMH is bilateral adrenalectomy. However, studies have demonstrated some evidence of biochemical and clinical improvement with unilateral adrenalectomy. A recent study by Osswald *et al.* compared 25 patients with BAMH who underwent unilateral adrenalectomy to 9 patients who underwent bilateral adrenalectomy and found that 84% of patients who underwent unilateral adrenalectomy had initial biochemical remission but only 32% of patients remained in biochemical remission at last follow-up with median follow-up of 50 months, compared to 100% in the bilateral adrenalectomy group. The rates of adrenal insufficiency were much lower in the unilateral adrenalectomy group (50% temporary, 5% permanent) compared to bilateral adrenalectomy (100%) for BAMH. Thus the authors conclude that in select patients with asymmetric hyperplasia or mild cortisol secretion it is reasonable to consider unilateral adrenalectomy as acceptance of mild hypercortisolism may be preferable to avoid lifelong glucocorticoid dependence and risk of adrenal crisis (30). The authors have performed unilateral adrenalectomy in BAMH with good results, always educating patients that a contralateral adrenalectomy may be necessary. How to choose which adrenal to remove first is purely based on the size of the adrenal gland and nodules, removing the larger gland first. In order to discern which adrenal gland is contributing the largest amount of cortisol, adrenal venous sampling has been performed (31), but to date results are disappointing. These complicated patients should be cared for in a multidisciplinary fashion in close collaboration with a pituitary/adrenal expert endocrinologist.

PPAD

PPAD is a rare cause of ACTH independent hypercortisolism occurring mainly in children and young

adults. Its name originates from the macroscopic appearance of the adrenal gland as it has characteristic small pigmented molecules in the cortex. The disease is associated with the Carney complex which is a complex of cardiac myxoma, pituitary adenoma, thyroid adenoma or carcinoma, testicular tumors, ovarian cysts, schwannomas and breast adenomas (32). On imaging the adrenals can appear normal or micronodules can be visible on one or both glands. The definitive treatment of choice to control hypercortisolism in patients with PPAD is bilateral MIS adrenalectomy (16).

ACC

ACC is a rare and aggressive primary adrenal malignancy (33,34). It is commonly diagnosed during evaluation of symptoms related to hormone excess as approximately 60% of ACC's release hormones, most frequently cortisol (20–30%) (35–37). Manifestations of hypercortisolism in ACC can be subclinical or overt and/or paired with androgen excess. Cortisol excess in ACC has been associated with decreased overall and recurrence free survival in retrospective studies (38). And as discussed above, it has also been associated with increased risk of postoperative complications due to the impact on the immune system, coagulation cascade and tissue healing (36).

The possibility of ACC should be considered in all patients with an indeterminate adrenal mass and hypercortisolism. While some patients will present with symptoms of malignancy such as weight loss, anorexia, cachexia or pain, a large proportion will not present with symptoms and therefore a review of preoperative imaging is essential to evaluate the risk of malignancy.

Imaging of an adrenal mass suspicious for ACC should include an adrenal protocol CT or MRI. On CT, ACC is usually large and heterogeneous with indistinct borders. Locally invasion of adjacent organs is common. Invasion of liver, kidney and vena cava is common on the right, whereas pancreas, spleen, renal vein and kidney may be involved on the left. On venous phase imaging vena cava tumor thrombus may also be identified, often originating at the level of the right adrenal vein and extending into the retrohepatic cava or extending from the left renal vein into the infrahepatic cava. If ACC is suspected, a CT of the chest should be performed to evaluate for pulmonary metastasis and identify any occult pulmonary emboli. FDG PET can also be used to help in the diagnosis and staging of patients with suspected ACC (18).

Complete surgical resection with negative margins is

the only curative option for ACC (33,39–42). Thus, the operative approach chosen must have the highest likelihood of achieving this goal. The AACE/AAES, the Society of American Gastrointestinal and Endoscopic Surgeons (SAGES) as well as the European Network for the Study of Adrenal Tumors (ENSAT) guidelines all agree that open adrenalectomy should be performed if ACC is suspected (43–45). These recommendations are due in part to the nature of ACC.

ACC is often a very soft and necrotic tumor; capsular disruption and fragmentation are easy to induce, particularly during direct tumor manipulation that commonly occurs in MIS. Furthermore, ACC tends to invade through the tumor capsule with microscopic disease present at the gland surface, thus minimizing direct contact with the tumor surface is essential to avoid violating the tumor capsule or causing disruption of disease at the surface of the gland (33). Thus, we agree with the aforementioned consensus guidelines and perform open adrenalectomy for known or suspected ACC.

Finally, the importance of surgeon experience in management of ACC cannot be overemphasized. Several studies have demonstrated higher rates of complete resection and improved outcomes when ACCs are managed by surgeons or at centers with expertise (40,46–49). Referral to high volume centers, experienced in treating patients with adrenal tumors and ACC, should be considered prior to proceeding with resection, regardless of operative approach.

The role of robotic adrenal surgery in management of patients with hypercortisolism and benign adrenal tumors

The first robotic adrenalectomy was reported by Horgan *et al.* in 2001 (50) and since that time has gained popularity due to the improved optics associated with 3D vision, the freedom of movement from the Endo-wrist instrumentation and improved surgeon ergonomics (51). While some argue that robotic technology is associated with higher cost and prolonged operative times (52) others have demonstrated that by limiting unnecessary instruments and energy devices as well as having an experienced surgical team the cost and operative times can be equivalent (51).

As previously mentioned, in patients with hypercortisolism, there is often increased central adiposity which can make traditional laparoscopic surgery quite challenging particularly when utilizing a retroperitoneal approach due to its inherently constrained space. The

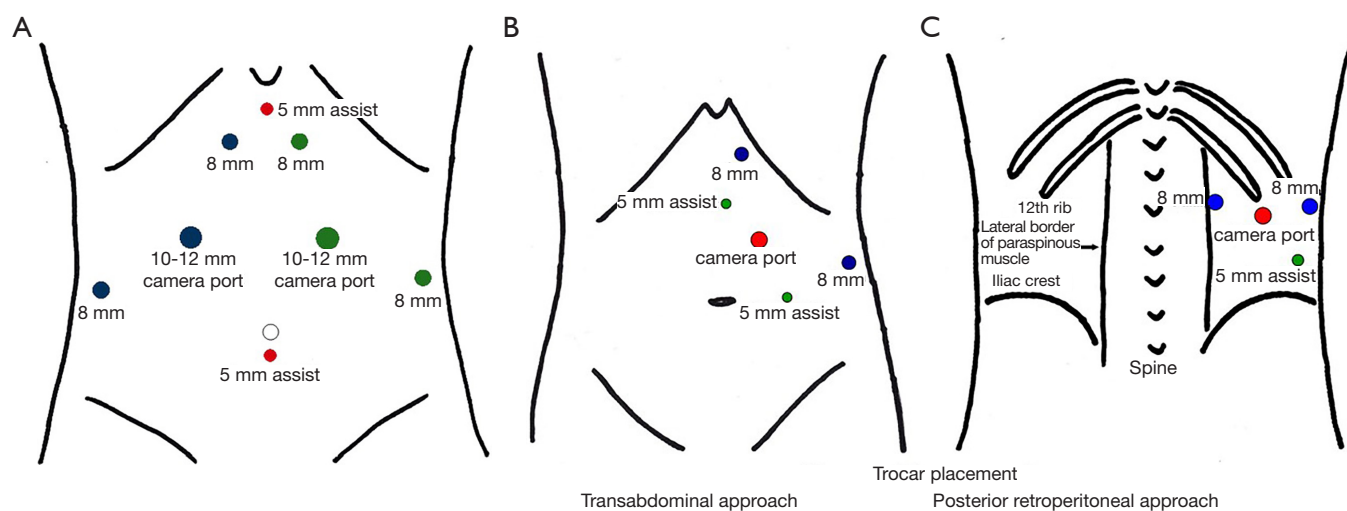


Figure 1 Suggested robotic adrenalectomy port placement for (A) bilateral transabdominal adrenalectomy, (B) unilateral (LEFT) transabdominal adrenalectomy and (C) unilateral (RIGHT) posterior retroperitoneoscopic adrenalectomy.

sharp 3D visibility, ability to reach and visualize tight places and the increased dexterity of the instruments helps the surgeon overcome space constraints both for retraction and dissection. It is the authors opinion that robotic adrenalectomy is easier to learn when compared to laparoscopic. For the already experienced adrenal team we did not observe a steep learning curve when our practice switched from laparoscopic to robotic systems to remove the adrenal gland 5 years ago; and in fact, operative times decreased over time specially when the Xi da Vinci system was introduced (53). We have observed trainees perform the steps of the RTA procedure with ease and efficiency when the robot is utilized (personal observation). Robot systems allow for easier handling of the rare complication of adrenal vein tear or injury to large veins as the surgeon can quickly repair the vessel with the added dexterity of the robotic system. Lastly, robotic systems allow the authors to address larger masses, those that sit on the renal vessels in the kidney hilum and/or occasional paragangliomas in difficult spaces that may not be easy to approach laparoscopically. In essence, we are able to “push the envelope” in our ability to remove the adrenal gland.

Robotic adrenalectomy operative technique

Patients are positioned in the lateral decubitus flank position on a bean bag for the transabdominal approach (TA) and the prone jackknife position on the Wilson frame for

PRA.

We acknowledge that there are many ways to place trocars in robotic procedures. Our recommended trocar placement for bilateral RTA (A), unilateral RTA (B) and unilateral RPR (C) adrenalectomy are depicted in *Figure 1*. The goals of trocar placement are to triangulate to the target gland, minimize collision of the robotic arms, and allow for a functional assistant trocars ports. The Xi platform may not require the same triangulation but our trocar placement suggestion works for both systems the Si and Xi.

Our approach for RPR and RTA has been described by our group previously (54) and are summarized step by step in *Table 3*.

Summary

The surgical management of hypercortisolism is often complex. In ACTH-dependent hypercortisolism, the first-line treatment focuses on the surgical removal of the site of ACTH production. Bilateral adrenalectomy should only be considered as a second line treatment for ACTH-dependent hypercortisolism, either CD or EAS and preferably after multidisciplinary discussion. ACTH-independent hypercortisolism is most commonly caused by a benign unilateral adrenal adenoma however consideration of much less frequently occurring etiologies of ACTH-independent hypercortisolism such as BAMH, PPAD and ACC is warranted in special populations to

Table 3 Step by step description of robotic adrenalectomy

Step	Right robotic posterior retroperitoneoscopic adrenalectomy	Left robotic transabdominal adrenalectomy
Step 1: entry	(I) 12 mm transverse incision below the tip of the 12 th rib; (II) Enter retroperitoneal space sharply with mayo scissors; (III) Develop space bluntly with surgeon finger	Enter abdomen using Optiview technique with 12 mm trocar in the mid-clavicular line, 5 cm inferior to costal margin
Step 2: trocar placement	(I) Place 8 mm robotic trocar lateral to initial incision; (II) Place 8 mm robotic trocar medial approximately 3 cm below junction of 12 th rib and paraspinous muscle; (III) Place 12 mm blunt balloon trocar in the access site; (IV) Additional (optional) 5 mm assistant port can be placed lateral and inferior to camera port	(I) Place two 8 mm robotic trocars in triangulation with camera port; (II) Dock the robotic (cadiere forceps left arm, robotic hook cautery right arm)
Step 3: create operative space	(I) Obtain “triangulation view” of medial muscle and blue layer of the peritoneum with blunt dissection; (II) Dock the robot (cadiere forceps left arm, robotic hook cautery right arm)	(I) Mobilize the splenic flexure of the colon; (II) Take down attachments between spleen and diaphragm and pancreas, kidney and adrenal allowing spleen and pancreas to be medialized
Step 4: Identification and mobilization of the gland	(I) Perform lateral to medial dissection detaching tissue above kidney; (II) Dissect adrenal gland and surrounding tissue from superior aspect of kidney	(I) Identify adrenal tissue and separate from pancreas superiorly; (II) Continue dissection medially and inferiorly
Step 5: identification and ligation of the adrenal vein	(I) Identify adrenal vein medially extending to vena cava; (II) Clip adrenal vein with 5 mm Hem-o-lock clips	(I) Identify adrenal vein medial and inferior draining in to renal vein; (II) Clip adrenal vein with 5 mm Hem-o-lock clips
Step 6: mobilization from the retroperitoneal fat	Remove adrenal gland from retroperitoneal attachments with hook cautery	Dissect adrenal gland with a rim of surrounding fat off of the retroperitoneum
Step 7: removal of gland	Place adrenal gland in bag and remove through 12 mm port	Place adrenal gland in bag and remove through 12 mm port

ensure appropriate surgical management. The majority of adrenalectomies performed for hypercortisolism can be performed utilizing a minimally invasive approach however if ACC is highly suspected open adrenalectomy should be performed. Robotic technology is a relatively new and useful technology when performing minimally invasive adrenalectomy. It can be utilized safely and cost effectively in experienced hands.

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Footnote

Conflicts of Interest: The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all

aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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