Selecting incision-dominant cases for robotic liver resection: towards outpatient hepatectomy with rapid recovery

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Background: The premise of minimally invasive surgery (MIS) is to minimize facial and muscle injury in order to enhance recovery from surgery. Robotic MIS surgery for resection of tumors in solid organs is gaining traction, though clear superiority of this approach is lacking and robotic surgery is more expensive. Our philosophy in robotically-assisted hepatectomy has been to employ this approach for cases where location of tumors make difficult a classical laparoscopic approach (superior/posterior tumors), and cases where the incision for an open operation dominates the course of recovery.

Methods: This is a retrospective review of a prospectively collected database.

Results: In this study we report 97 cases of liver resection subjected to the robotic approach, of which 90% were resected robotically. The mean operative time was 186 ± 9 min; mean blood loss was 111 ± 15 mL, and complications occurred in 9%. Two thirds of the patients remained in hospital 3 days or less, including three patients subjected to hemihepatectomy (2 left and 1 right). Fourteen individuals were discharged on the same day. The strongest predictors of long hospital stay (>3 days) were major hepatectomy (P=0.007), complications (P=0.008), and operative time >210 min (P=0.001).

Conclusions: With thoughtful case selection, this is a first demonstration that hepatectomy can be conducted as an out-patient or short-stay procedure.

Keywords: Adoption of minimally invasive surgery (MIS); enhanced recovery after surgery (ERAS); robotic MIS hepatectomy; surgical outcome

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Introduction

Liver resection has evolved greatly over the last three decades. Once an operation with very high risk of hemorrhage and high mortality rate (1), most hepatectomies are now performed at major centers with a mortality rate routinely less than 3% (2). Transfusion is required in only a minority of patients (3). Techniques of liver resection have also evolved greatly. Minimally invasive hepatectomy was once discouraged. Recent advancements in equipment,

technique, and training have resulted in general acceptance of laparoscopic hepatectomy (4). A number of studies have shown a trend for earlier discharge and adequate oncologic outcomes with laparoscopic hepatectomy (4), though clear superiority of the technique in objective outcome is sparse because of difficulties in measuring recovery outcomes in general.

Robotic hepatectomy is the latest evolution of minimally invasive surgery (MIS). With 3D optics, wristed instruments, integrated stapling, and sealing instruments, many tasks that were technical challenges for laparoscopic surgery are overcome including suturing and fine vascular dissection. These technical advantages have come at a higher monetary cost (5,6), sufficient to demand a justification by data on outcome to show advantage for patients. Most early robotic hepatectomy papers (7,8) had a high percentage of major liver resections (>35%), and touted as advantage of robotic hepatectomy the low conversions rate. In one paper (7), the advantage was cited as "more likely to complete operation in totally minimally invasive approach".

In the current paper we proposed a different philosophy for robotic hepatectomy. We propose that the best operations to be performed robotically are those involving removal of small amounts of parenchyma, particularly for lesions placed in parts of the liver hard to reach by laparoscopy. For these poorly placed tumors, usually a large incision is required to resect tumor. We propose that it is for these "incision-dominant" operations, where the facial and muscle incision and not the physiology of liver regeneration dominate recovery, that robotic hepatectomy is most likely to help patients and be measured to do so. Following is our preliminary series demonstrating many of the advantages of such an approach.

Methods

Patients

This is an observational study of a prospectively maintained liver resection database. We included all patients 18 years or older, who underwent exploration with intent to perform a robotically-assisted liver resection. This paper includes patients submitted to robotic resection by the senior author (Yuman Fong) at the Memorial Sloan-Kettering Cancer Center, and operations performed by all surgical authors at the City of Hope Medical Center (Yuman Fong, Susanne Warner, Laleh Melstrom, Gagandeep Singh, Byrne Lee, Yanghee Woo). We excluded patients undergoing liver biopsy. The study was approved by the City of Hope Institutional Review Board #15097.

Data for these patients were extracted from the database and additional data relevant to the study were extracted from the electronic medical records. Data examined included demographics (age, gender, body mass index, American Society of Anesthesiologists classification, Eastern Cooperative Oncology Group performance status, comorbidities), indications and perioperative variables (extent of liver resection, estimated blood loss, duration of surgery, concomitant procedures). The size of liver resection was defined using the International Hepato-Pancreato-Biliary Association Couinaud's classification (9). Major hepatectomy was thus defined as a resection of 3 or more segments. Margins were defined as positive (<1 mm), close (1–10 mm), and negative (>10 mm).

Complications and readmissions

All complications during the index hospitalization and readmission were recorded up to 90 days after surgery. Mortality was recorded at 90 days. Readmissions within 30 days after discharge from index hospitalization per the CMS definition were recorded. Each complication was categorized according to the Clavien-Dindo complication grading system as detailed elsewhere (10). Briefly, complications were graded from 1 to 5, where 1 and 2 usually indicate management with either oral or intravenous pharmacologic treatment; and where grade 3 indicates procedures requiring involvement of interventional radiology or gastroenterology, or return to the operating room. Grade 4 indicates treatment in the intensive care unit for organ dysfunction or permanent disability as sequelae of the complication; grade 5 indicates patient death.

Robotic operations

Patients were evaluated for suitability of the robotic approach depending on the location of their tumors, the quality of their liver, and the patient's clinical status. Contraindications for robotic resection were invasion of the inferior vena cava (IVC) or base of hepatic veins close to the IVC, invasion of the main right or left portal vein, and need for vascular reconstruction or bile duct resection.

The Si da Vinci Surgical System or the Xi da Vinci Surgical System was used with four arms. Operative procedure was as described before (11). Patients were secured to a table with a footboard that allowed for 30 degrees of reverse trendelenburg. For the Si system, a 12-mm trocar was used to place a camera and three 8-mm ports were used for the instrument arms. Liver parenchyma was divided with a combination of harmonic scalpel, the vessel sealer, the PK bipolar dissector, or the stapler. The patient cart was always placed by the patient's feet. The scrub nurse was positioned by the patient's right leg, and the assistant surgeon or surgical physician assistant was positioned by the patients left leg.

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Variable	All	Robotic completion	
Ν	97	87	
Age, median [range]	62 [40–91]	62 [40–91]	
Male sex	52 (54%)	44 (51%)	
Operative time (mean \pm SEM)	197±6 min	186±9 min	
Blood loss (mean ± SEM)	144±15 cc	111±15 cc	
Complications	10 (10%)	8 (9%)	
Operations			
Major resection	13 (13%)	8 (9%)	
Minor	84 (87%)	79 (91%)	
Segments 3, 4B, 5, 6	51 (53%)	50 (57%)	
Segments 1, 2, 4A, 7, 8	33 (34%)	29 (33%)	

Table 1 Demographics of study population

Statistical analysis

Bivariate analysis of categorical variables was performed using the Chi-square test. Criterion for statistical significance set at P<0.05.

Results

Demographics

The study population consisted of 97 patients subjected to exploration for possible robotically assisted hepatectomy. Median age was 62 years with range 40–91 (*Table 1*). Fifty four percent were male. The diagnosis associated with these operations included metastatic tumors to the liver (65%), primary hepatocellular carcinoma (10%), biliary cancers (11%), and benign tumors and conditions (14%).

Conversion to open surgical procedure occurred in ten patients, mostly due to more extensive disease found. For the 87 procedures completed robotically, the mean operative time was 186±9 (median 170, range, 30–486) min. Mean blood loss was 111±15 (median 50, range, 25–800) mL. Complications occurred in 8 individuals (9%), and included two cases of pneumonia, two intra-abdominal abscesses, ileus, cellulitis, and urinary retention. There were no perioperative deaths.

Procedures performed included eight major hepatectomies: two right hepatic lobectomies (one with in continuity diaphragmatic resection, *Figure 1*), and six left hepatic lobectomies. The majority of procedures were minor hepatectomies (91%), 57% of which involved the lower segments of liver (3, 4B, 5, 6), and 33% involving superior segments (1, 2, 4A, 7, 8). Minor procedures included 13 left lateral sectorectomies, and seven right posterior sectorectomies.

Hospital stay

Two thirds of the patients remained in hospital 3 days or less, including three patients subjected to hemihepatectomy (2 left and 1 right) (*Figure 2*). The three patients subjected to minor hepatectomies who stayed beyond 1 week in hospital were all over 80 years of age and stayed due to placement issues.

The strongest predictors of long hospital stay (>3 days) were major hepatectomy, complications, operative time >210 min, and blood loss >100 cc (*Table 2*).

Fitness Sensor Tracking

For the patient subjected to right hepatic lobectomy and incontinuity diaphragmatic resection (*Figure 1*), we tracked her preoperative activity and post-operative recovery with a fitness tracker (Vivofit[®], Garmin Ltd., Schaffhausen, Switzerland). The activity curve is shown in *Figure 1D*. Her level of activity was approximately 6,000 steps prior to surgery and recovered to 6,000 steps by day 4. She was discharged on day 2 post-operatively.

Discussion

Recovery from hepatectomy includes overcoming (I) surgical stress, (II) systemic effects of liver regeneration, (III) incisional discomfort, and (IV) associated ileus. MIS, including robotically-assisted MIS surgery is intended mainly to alleviate discomfort associated with a large incision and decrease deleterious effects associated with evaporate losses during a laparotomy. The current paper suggests that the patient most likely to benefit from robotically-assisted MIS hepatectomy is that patient subjected to an incision-dominant procedure. In particular, for minor hepatectomies in regions of the liver poorly accessible to laparoscopy (4), that normally would require large open incisions, robotic MIS hepatectomy offers an attractive solution with rapid recovery.

Robotic MIS hepatectomy as a field is moving past its infancy. Nota *et al.* (12) recently reported a collected series analysis of the 12 best studies up to 2015 (7,8,13-22).

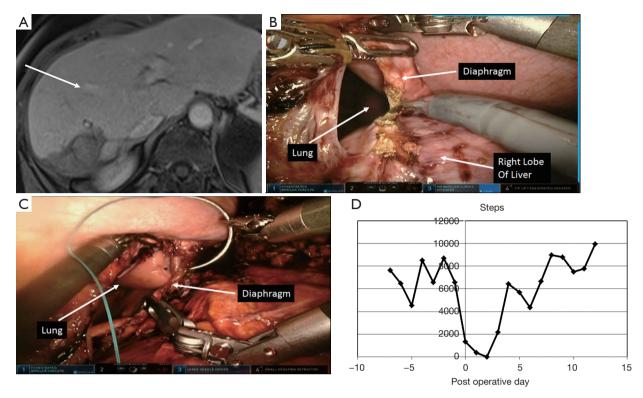


Figure 1 Representative patient undergoing robotic right hepatectomy. (A) CT scan of a 42-year-old patient with metastatic cancer in segment 6, 7 and 8 of liver. Involvement of the junction of the right anterior and posterior pedicles resulted in compensatory hypertrophy of the left liver. Note large left liver as demarcated by the middle hepatic vein (arrow); (B,C) this patient also has involvement of the diaphragm requiring in-continuity resection of the right diaphragm and primary repair; (D) the data form the Vivofit fitness monitoring device. Note that the patient was back to 6,000 steps at Day 4 post-operatively.

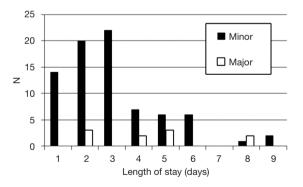


Figure 2 Length of hospital stay for robotic hepatectomy according to minor or major (≥3 segments).

There is no doubt that hemi-hepatectomies can be performed with safety by the robotically-assisted approach (7,8). However, for patients requiring major hepatectomy, particularly operations that are long and requiring abundant fluid administration and blood transfusion, the recovery is likely to be dominated by liver regeneration or by surgical trauma. Consequently, operative times are long and hospital stays are very similar to that of open hepatectomy (*Table 3*). In these operations where recovery is dominated by the liver regenerative process, it will be difficult to prove superiority of a robotic approach, especially since the robotic approach will at least in the short term be more costly (5,6). Major hepatectomy will remain an operation performed by experts. The choice of whether to perform these robotically will be according to philosophies of individual centers. We have chosen to be highly selective in robotic hemi-hepatectomies, restricting these operations to cases where early discharge is likely: cases involving young, fit, motivated patients, with tumors away from the hilus or confluence of the hepatic veins.

The published data on robotically-assisted hepatectomy should be segregated into three groups: (I) major hepatectomy, (II) minor inferior hepatectomy, and (III) minor superior/posterior hepatectomy (*Table 3*). Minor

Table 2 Predictors of length of stay (LOS)								
Parameter	LOS ≤3 days	LOS >3 days	Р					
All	58	29						
Gender			0.9					
Male	29	15						
Female	29	14						
Size of resection			0.003					
Minor	56	22						
Major	2	7						
Complications			0.009					
No	56	23						
Yes	2	6						
Operative time (mi	n)		0.001					
≤210	49	15						
>210	9	14						
Blood loss (mL)			0.09					
≤100	49	20						
>100	9	9						

Table 2 Predictors of length of stay (LOS)

inferior hepatectomy (segment 3, 4B, 5, 6, or left lateral sectorectomies) are procedures also suited for traditional laparoscopic surgery (4), since these are regions of the liver approachable by straight laparoscopic instruments. At present, either a laparoscopic or a robotic approach can be used to deliver the benefits of MIS surgery for patients with tumors in these parts of the liver (26), and traditional laparoscopic approach is less expensive (5,6). We predict however, that as 3D, high definition laparoscopy, and other high priced technology gain penetrance in laparoscopy, and as other robotic platforms reach market, routine laparoscopy and robotically-assisted MIS will approach price equivalency. The robotic approach has also proven to provide a low barrier for adoption of MIS surgery. As availability of robots increases worldwide, and as cost of robotic operations decrease, it is more likely that future surgeons and current surgeons dedicated to open operations will more likely learn the robotic approach than routine laparoscopic.

Minor superior-posterior hepatectomies (segment 1, 2, 4A, 7, 8, or right posterior sectorectomies) consist of operations thought poorly approachable by traditional laparoscopy (4). This is because it is difficult to reach these

areas and performed skilled moves with straight instruments. Thus, tumors in these parts of the liver currently are approached by large open incisions, even when the tumors are small. If a randomized trial is desired for operations in superior, posterior segments, the randomization has to be against open operation. Other investigators have also recognized the advantages of a robotic approach for tumors in these regions (12,29). The articulated instruments and the 3D optics of the surgical robot are well suited for operations in these anatomically "remote" areas of the liver. The current study confirms that hepatectomy for tumors in these regions of the liver to be ideal for robotic MIS approach, achieving very low complication rates and very short hospital stavs.

Measures of quality of hepatectomy and major surgery in general have traditionally relied on mortality and complication rates. In recent years, the mortality from hepatectomy at major center have approached 0-1% (31), and complication rate for minor hepatectomies are now routinely less than 20%. Surrogate markers for quality, such as blood loss, ICU stay, hospital stay, or re-admission to hospital are now used to stratify quality of surgical program. However, with improving surgical and anesthesia techniques, hemorrhage is a rare problem, and many "bloodless" hepatectomy (32) series are being reported with little use for transfusion. In the current study we show that good patient selection can lead to low blood loss, low complication and short stay for robotically assisted hepatectomy.

Ultimately, MIS surgery should be measured by functional recovery, since the biggest difference from open surgery is the size of the muscle and facial incision. Many have attempted to use (33) quality of life instruments to document the superiority of the MIS approach, but have been hindered by the how cumbersome some instruments are and the limited number of time points that can be accessed. We have begun a program evaluating use of athletic fitness devices to measure functional recovery (*Figure 1D*). We recently presented our pilot study demonstrating feasibility and acceptability of this approach (34). We are now focusing a study of such post-operative fitness tracking for MIS surgery.

Until recently, the idea that hepatectomy could in a major subset of patients be an outpatient procedure would have been unthinkable. Not long ago, papers reporting hepatectomy outcomes talked about length of ICU stay, and hospital stays of two weeks were routine. The current paper is one of the first reporting feasibility of outpatient

Study	n*	Operating time, mean± SD or median [range], min	Conversion (%)	Positive surgical margin (%)	Blood loss, mean ± SD or median [range], mL	Length of stay, mean ± SD or median [range], days	Complication (%)	Mortality
Resections of >4 segme	nts							
Nota <i>et al.</i> 2016* (12)	99	405±100	8	0	380±505	11±6	26	0
Chen <i>et al.</i> 2016 (23)	34	343 [140–715]	0	0	182	8.9	6	0
Lai <i>et al.</i> 2016 (24)	100	207±77	4	4	335 [5-3,500]	7.3±5.3	14	0
Quijano <i>et al.</i> 2016 (25)	20	282	5	0	n/a	13.4	20	0
Minor resections (2, 3, 4	B, 5, 6)							
Nota et al. 2016* (12)	81	215±65	1	2	230±310	5±2	19	0
Kim <i>et al.</i> 2016 (26)	12	404±139	n/a	0	225 [125–275]	7 [7–8]	25	0
Salloum <i>et al.</i> 2016 (27)	16	190±7	13	0	247±239	6±4	13	0
Lee et al. 2016 (28)	70	252 [97–620]	6	2	100 [2–2,500]	5 [2–22]	11	0
Present series	50	166±59	0	0	96±140	2.9±2.0, 3 [1–9]	12	0
Minor resections (1, 4A,	7, 8)							
Nota et al. 2016* (12)	17	220±60	0	2	170±120	5±1	24	0
Montalti <i>et al.</i> 2016 (29)	36	306±182	14	11	415±414	6±2.9	19	1 (2.8%)
Boggi <i>et al.</i> 2015 (30)	12	260 [115–430]	8	0	253 (50–600)	8.5 [7–96]	33	0
Present series	29	191±75	18	0	125±159	3.2±1.5, 3 [1-6]	7	0

Table 3 Past studies for robotic hepatectomy

*, collected series. n/a, not applicable.

or short stay hepatectomy. The results of the current paper indicate that selecting patients who are medically fit and who have physiologically favorable lesions (small parenchymal resection) and performing a short operation (to minimize surgical stress), and with small fascial and muscular incisions, can result in outpatients or short stay hepatectomy. We have also recently published our enhanced recovery after surgery (ERAS) pathways for hepatectomy, which includes a pathway for robotically-assisted hepatectomy (35).

Conclusions

Surgical outcomes are optimized when the surgeon (I) selects patients for the surgical approach most beneficial, and (II) performs the operation with skill. Selecting incision

dominant procedures ensures highest likelihood for benefit from a robotic MIS approach. Robotically-assisted MIS surgery brings advanced tools to complement the skills of hepatic surgeons to further enhance outcomes in these difficult operations.

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Footnote

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

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Ethical Statement: The study was approved by the Institutional Review Board of City of Hope (No. 15097).

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