

Robotic surgical system completed radical gastrectomy for gastric cancer after liver transplantation: case report and systematic review

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Abstract: *De novo* gastric cancer (GC) is uncommon in clinic after liver transplantation. Until now, only a few cases have been reported, and radical resection combined with comprehensive treatment remains the major therapeutic method. Two patients with De novo GC after liver transplantation were successfully treated with D2 lymphadenectomy using da Vinci robotic surgery system at our Hospital. Both operations were successful, and the duration of case 1 and 2 was 315 and 275 minutes respectively, with the estimated blood loss of 145 and 125 mL. The patients were discharged on day 7 and 9 after surgery, and no complications occurred. Postoperative pathological stages were pT4aN3aM0, stage III B and pT4aN2M0, stage III A respectively. Case 2 developed gastric retention 3 weeks after operation, and gastroscopy revealed edema within anastomotic stoma. Conservative treatment using feeding jejunal tube was performed. After 2 weeks, normal diet was gradually restored. Patients were in good health without long-term complications. Therefore, GC surgery using Da Vinci robot surgery system after liver transplantation is safe and feasible.

Keywords: Da Vinci robot surgery system; gastric cancer (GC); gastrectomy; liver transplantation; case report

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Introduction

Da Vinci robot surgery system has been used in the surgery of gastric cancer (GC). The operations were performed on GC patients at early and advanced stages. The surgical methods have been developed from distal to proximal radical gastrectomy, and even in radical total gastrectomy and gastric remnant after partial gastrectomy (1,2). New cases of GC after liver transplantation are rarely reported, and no robotic surgery has been reported so far. Only a few cases have been reported, and radical resection combined with comprehensive treatment remains the major therapeutic method. One case of laparoscopic-assisted radical resection has been reported in Korea. In this study, with the assistance of robotic GC surgery (3), radical GC surgery was successfully performed on two patients after liver transplantation at Southwest Hospital on October 23 and December 25, 2018. Follow-up examination was carried out 6 and 4 months after surgery, respectively, and the therapeutic outcome was good. Written informed consent was obtained from each patient prior to operation, and the information of patients was anonymous. We present the following case in accordance with the CARE reporting (available at http://dx.doi.org/10.21037/tcr-19-1786).

Case presentation

Case 1, male, 63 years old, was diagnosed with primary liver cancer and received orthotopic liver transplantation at Southwest Hospital in 2007. During the 11 years after

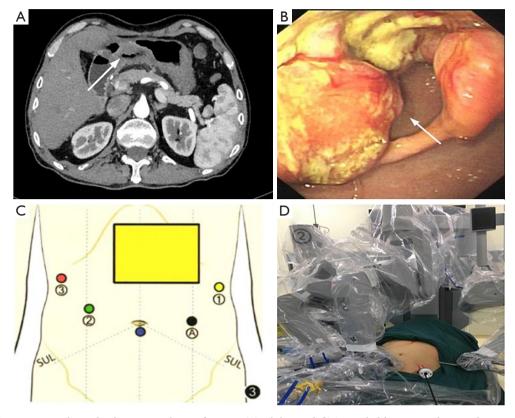


Figure 1 The diagnostic results and robot manipulator of case 2. (A) Abdominal CT revealed lesions in the antral region; (B) gastroscopy indicated lesions in case 2; (C) layout of robot manipulator point card; (D) solid assembly of robot manipulator.

surgery, the quality of his life is improved. No medical or family history of liver cancer and GC were presented. However, the patient was hospitalized and subjected to enhanced CT scan in September 2018 due to pain or discomfort centered in the upper abdomen. The results revealed irregular shape of his gastric antrum wall, and the mucosa was rigid and rough, small curvature of stomach and pylorus were scattered with enlarged lymphnodeshadows, up to $\sim 4.0 \times 2.3$ cm² in size. No recurrence and distant metastasis of hepatocellular carcinoma was detected. Gastroscopy revealed giant irregular ulcer in antrum. Additionally, the pathological diagnosis was adenocarcinoma of antrum. Physical examination revealed that the body mass index was 25.08 kg/m², no abdominal mass or enlargement of Virchow lymph nodes was found, and there was an inverted L-shaped incision (~50 cm) in the right upper abdomen. Diagnostic results: (I) adenocarcinoma of gastric antrum; (II) postoperative complication of liver transplantation; (III) varicose veins in the upper digestive tract.

Case 2, a 64-year-old male, underwent orthotopic liver

transplantation at Southwest Hospital in January 2018 due to acute-on-chronic liver failure. The quality of his life was improved in the following 11 months. No medical or family history of liver cancer and GC were presented. The patient was hospitalized and subjected to CT scan in September 2018 due to upper abdominal discomfort. The results indicated that the gastric wall in lower gastric body and antrum were unevenly thickened, and soft tissue density was increased. Mucosal disruption was also detected in the lesion area, where large ulcer (~5.0 cm in diameter) could be observed. The gastric wall of lesion area was stiff, and lesion boundary was unclear. In addition, the serosal surface was not smooth, and the lesion extended to pylorus, consequently resulting in pyloric stenosis (Figure 1A). Gastroscopy revealed giant irregular ulcer in antrum (Figure 1B). The pathological diagnosis was poorly differentiated adenocarcinoma in antrum. Physical examination revealed that the body mass index was 20.37 kg/m^2 , no enlargement was detected in supraclavicular lymph node, and there was an inverted L-shaped incision (~50 cm)

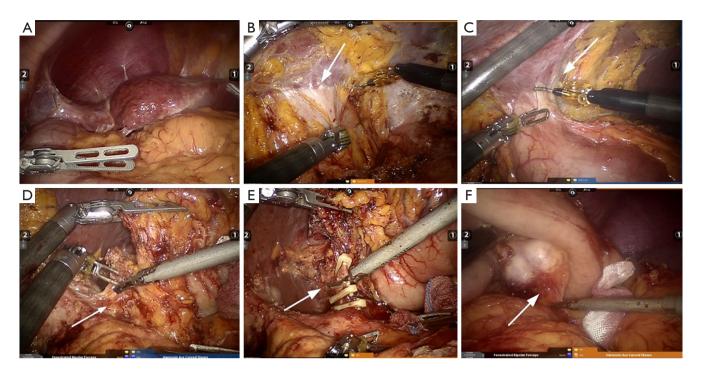


Figure 2 Representative images of the operation. (A) During the operation, omentum and hepatic flexure of colon moved upward to enclose the first hepatic hilum and adhered to the hepatic margin; (B) the hepatic flexure of colon adhered to the right abdominal wall; (C) in case 2, anterior wall of gastric antrum adhered to the upper abdominal wall; (D) in case 2, lymph nodes were scanned; (E) in case 1, after bioclips were placed, ultrasound scalpel was applied; (F) in case 1, the posterior wall of stomach was turned upward during the operation, and the pancreatic-gastric ligament was exposed. The enlarged lymph nodes were cleared.

in the right upper abdomen. Diagnostic results: (I) adenocarcinoma of gastric antrum; (II) postoperative complication of liver transplantation; (III) type II diabetes mellitus.

Radical subtotal gastrectomy with robotic Da Vinci system was planned for both patients.

Surgical procedure

Tracheal intubation and intravenous anesthesia were performed. The supine position was used, and the legs were separated. The abdominal wall was punctured using 5-hole method: 12 mm Trocar was inserted as the lens hole 2 cm from the lower umbilical margin, and 8 mm Trocar was placed beneath the left axillary front rib and used as the main manipulation hole of robot-arm no.1. Additionally, 12 mm Trocar was inserted 1 cm under the left clavicle midline as the assistant manipulation hole, and 8 mm Trocar was placed 3 cm beneath the right clavicle midline was with as the manipulation hole of robot-arm no.2, Furthermore, 8 mm Trocar was inserted under the right axillary front rib as the manipulation hole of robot-arm no.3. The interval between Trocars was ≥ 8 cm (*Figure 1C,D*). No ascites was found, and no metastasis was detected in peritoneum, omentum, pancreas, spleen, colon and pelvic. Both patients received orthotopic liver transplantation, and the omentum moved upward to encapsulate the porta hepatis and adhered to the abdominal wall (Figure 2A). In case 2, the left gastroepiploic membrane partially adhered to the splenic flexure and the left upper abdominal wall, and the anterior gastric wall also adhered to the abdominal wall. For the installation of Da Vinci robot system, the assembly of robot-arm no.1, no.2 and auxiliary holes was completed. The bipolar electrocoagulation of robot-arm no.2 was alternately coordinated with the hook of no.1. When the adhesion between omentum and the left upper abdominal wall was separated, the robot-arm no.3 was assembled, and the pyloric region of the antrum was exposed. The tumors were located prepyloric region of the stomach in both patients. The sizes of tumors were $\sim 3 \times 3$ and $\sim 4 \times 5$ cm² in case 1 and 2 respectively, which had penetrated the serosal surface. Furthermore, the adhesion in hepatic hilum, gastric

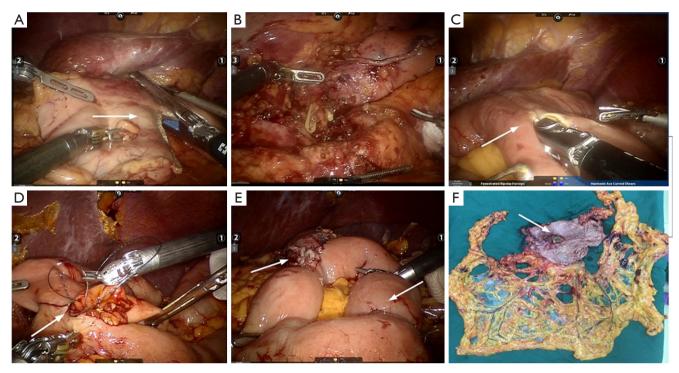


Figure 3 Representative images of the operation. (A) Distal gastrectomy was performed; (B) the stomach bed was cleared; (C) jejunostomy was carried out; (D) continuous suture was used for closure; (E) the digestive tract was reconstructed by gastrointestinal and intestinal anastomosis; (F) in Case 2, surgical specimens revealed large ulcer lesions on the lesser curvature side.

anterior wall and spleen was isolated. In addition, ascending colon, duodenum and hepatic hilum were adhered, which was completely separated using the ultrasound knife attached to robot-arm no.1 (*Figure 2B,C*).

The greater omentum was lifted, and the transverse colon was separated using ultrasonic scalpel. The left gastroepiploic artery and vein were isolated, and the lymph nodes of group 4sb were dissected. Subsequently, the right gastroepiploic vessels were isolated, and the lymph nodes of group 4d, 14v and 6 were removed. Furthermore, the duodenal bulb was exposed and the duodenum was dissected using an intraluminal closure device, and stomach was rotated to the left. The common bile duct, portal vein and proper hepatic artery were encapsulated by fibrous tissues. As a result of liver transplantation, right gastric artery or vein was not observed. Lymphatic adipose tissue was dissected in group 5 and 12a. Lymph nodes of group 8a were removed along the surface of common hepatic artery at the upper edge of the pancreas (*Figure 2D*).

In case 1, the left gastric artery was not found, but compensatory hypertrophic arteries and blood supply in the lesser curvature from the accompanying veins were observed. After bioclips were placed, ultrasound scalpel was applied (*Figure 2E*). The lesser curvature was isolated along the lesser curvature side of the stomach. The lymph nodes in group 3 were notably enlarged, (~3 cm in diameter), which were subsequently removed (*Figure 2F*). In case 2, the left gastric artery was exposed, lymph nodes and adipose tissue of group 7, 9 and proximal splenic artery (group 11p) were dissected, and left gastric artery and vein were clamped with two Tyco bioclips and cut with ultrasound scalpel.

The distal gastrectomy was performed using a 60 mm cutting closure device at 1 cm beneath the lesser curvature side of antrum and above the 1/3 boundary of the greater curvature side (*Figure 3A*). The distal gastrectomy and isolation of D2 lymph nodes were completed (*Figure 3B*). The reconstruction of digestive tract was completed using type II + Braun anastomosis with the assistance of Da Vinci Si robotic surgical system. Disposable laparoscopic stapler was used to perform anterior gastrojejunal side-to-side and jejunojejunal anastomosis. The continuous closure was performed using 3-0 barbed suture (*Figure 3C*,*D*,*E*). Indwelling drainage was completed, and an incision (~3 cm) was made at umbilical Trocar to remove the specimen (*Figure 3F*).

Results

Distal subtotal gastrectomy + D2 radical lymph node dissection were successfully performed on both patients.

Case 1

The operation last for 315 minutes, and the bleeding volume was 145 mL. The urinary catheter and nasogastric tube were removed 46 and 70 hours after surgery respectively, the anal exhaust was observed 48 hours postoperatively. Semi-liquid diet was taken by the patient orally 3 days after operation, since then, immunosuppressant mofetil (1,000 mg, BID) and tacrolimus (10 mg, BID) were given. The abdominal drainage tube was removed 6 days after surgery, and the patient was discharged on day 7. No complications such as abdominal hemorrhage, wound infection, anastomotic leakage and stenosis were observed. The pathological diagnosis was medium differentiated adenocarcinoma of antrum. A total of 19 lymph nodes were analyzed and confirmed by pathologist, and metastasis was found in 11 lymph nodes. The postoperative diagnosis was moderately differentiated gastric antrum adenocarcinoma pT4aN3aM0, stage III B. The first chemotherapy was performed 3 weeks after operation using SOX, and 8 cycles of treatment were planned. Intravenous chemotherapy was carried out for three times. At present, follow-up is still in progress. During the follow-up period, the patient is generally in good health and no complications occurred.

Case 2

The operation time was 275 minutes and the bleeding volume was 125 mL. The urethra and gastric tube were removed 36 and 70 hours after surgery respectively. The anal exhaust was observed 48 hours postoperatively. Semiliquid diet was taken by the patient orally 3 days after operation, and the treatment using mofetil (1,000 mg, BID) and tacrolimus (10 mg, BID) were resumed. The abdominal drainage tube was removed 8 days after surgery, and the patient was discharged on day 9. No postoperative complications occurred. The pathological diagnosis was poorly differentiated adenocarcinoma; 21 lymph nodes were checked by pathologist, and metastasis was detected in 2 of them. The postoperative diagnosis was pT4aN2M0, stage III A of moderately differentiated adenocarcinoma of the gastric antrum. Gastric retention was conducted since 3 weeks post-operation. Gastroscopy revealed edema in

anastomotic stoma. Conservative treatment was performed using jejunal tube. After 2 weeks, normal diet was gradually restored. Chemotherapy was initiated 2 months after operation using SOX. During the follow-up period, the patient is generally in good health, but suffers from acid reflux and belching occasionally.

Anesthesia and posture were the same as robotic surgery for primary GC. In both cases, an inverted L-shaped incision (~50 cm) in the right upper abdomen was used for liver transplantation. Due to local adhesion in the right upper abdomen, pneumoperitoneum was established and the first Trocar was placed 2 cm from the bottom edge of umbilical fossa to avoid damage to the intestinal tract. The remaining 4 Trocars were punctured under visual guidance. Firstly, the puncture of the left upper abdomen was completed, and an arm was fitted with an electric hook. Subsequently, Trocar of the second and third arms on the right side of the Da Vinci rob were assembled. End-gripping forceps were used for robot-arm no.3, which allowed larger contact area with intestinal canal and dispersed the pressing force evenly without damaging intestinal wall. They can also provoke liver exposure and help to reveal visual field, thus the efficacy was improved. The 2nd arm coordinated with the 1st arm for the fine-tuned movement during the operation, which provided more flexibility. Both arms were equipped with sharp and compact grippers. The tension between the 2nd arm and the tissues isolated by the 3rd arm was appropriate, which makes it possible for the electric hook and ultrasonic scalpel on the 1st arm to operate precisely.

The estimated blood loss in two cases was 145 and 125 mL, and the patients were discharged on day 7 and 9, respectively. Both patients benefit from the minimally invasive surgery. The second patient developed mild anastomotic edema 3 weeks following operation, but was recovered 2 weeks after conservative treatment.

Discussion

The most common types of cancer occurred after liver transplantation are non-melanoma skin cancer, respiratory and digestive system tumors, as well as post-transplant lymphoproliferative disorders (4). Immunosuppressive agents could be a risk factor due to the long-term suppression of immune system, which makes it easier for the mutated tumors to escape the immune surveillance, consequently leading to the formation of malignant tumors (5). In addition, age and consumption of alcohol and tobacco

Table 1 The cases of gastric cancer after liver transplantation

Authors	Year of publication	Country	Age/sex	LDLT malignancy	Treatment	Stage	Chemo-therapy
Nagata et al. (15)	2007	Japan	57/F	24 month	Open	_	None
Arslan <i>et al</i> . (16)	2011	Turkey	51/M	120 month	Open	IB	None
Lee et al. (13)	2011	Korea	72/M	180 month	LAP	IA	None
Shimizu <i>et al</i> . (8)	2012	USA	60/M	30 month	Open	IIA	None
Li <i>et al</i> . (17)	2013	China	55/M	108 month	Open	IV	None
Takehara et al. (18)	2014	Japan	65/F	96 month	Open	IIIA	S-1
Xiao et al. (14)	2016	China	41/M	144 month	Open	IIIA	_
Zhen <i>et al</i> . (19)	2016	China	68/F; 67/M	43/57 month	Open	-	None/YES
Yang et al. (11)	2016	China	63/M	96 month	Open	IIIC	None
Zhang et al. (20)	2017	China	64/F	120 month	_	П	PF
Present 2 cases	2018	China	63/M;64/M	132/11 month	Robtic	IIIB/IIIA	SOX

may be associated with newly formed malignant tumors after liver transplantation (6,7). Solid tumors are the main cause of liver transplantation-related mortality. It has been reported that the incidence of GC in liver transplant recipients is also increased, especially in Asian, where liver disease and GC pose great threat to public health (8-11).

Radical resection remains the major treatment for GC after liver transplantation. Gong et al. (12) reported that 19 of 2968 liver transplantation recipients were diagnosed with GC, and ESD was performed on 10 patients. Surgical resection was carried out on the remaining 8 patients as an initial treatment. One patient received chemotherapy first due to the peritoneal seeding revealed by preoperative CT scan. Until now, only a few cases of GC after liver transplantation have been reported (11,13,14). As presented in Table 1, Lee et al. (13) conducted an operation of laparoscopic-assisted radical gastrectomy for GC in 2011, and the others reported that laparotomy was performed. Compared with the surgery of primary GC, the operation of GC after liver transplantation is more difficult. The main reason is that the anatomical structure of upper abdomen has remarkably changed after liver transplantation: (I) extensive adhesions are formed among the organs and between the organs and abdominal wall; (II) the blood vessels in the hilum of liver are disrupted including the right gastric artery, which is an essential blood vessel in the stomach. In addition, the common hepatic artery becomes local dense adhesion after losing its normal sheath; (III) due to the exposure of blood vessels in the hilum during liver transplantation, the lymphatic reflux in the suprapubic region is impaired, especially the lymphatic reflux was blocked in group 5, 8 and 12.

Compared with traditional laparotomy and laparoscopic surgery, robotic surgery system provides better visual field exposure/clarity and operation flexibility. It also allows us to perform the operation in narrow space, and improves the relaxation of abdominal adhesion (3,21-23). In 2003, Hashizume *et al.* (24) reported the first successful robotic GC surgery. Since 2010, our team has been focused on robot-assisted radical gastrectomy for GC. At present, >1,000 operations have been performed, including 20 cases on gastric remnant after partial gastrectomy. In this study, robotic surgical system was used to perform radical gastrectomy on two patients.

During the operation, it is difficult to separate the adhesion of porta hepatis precisely without damaging the liver surface, ascending colon and duodenum. Attention should be paid to protect the common bile duct and portal vein during the precise isolation using ultrasonic knife and electric hook. No anatomical alterations were observed in the fundus and greater curvature of stomach, so the dissection of left gastroepiploic vessels and lymph nodes in group 4sb were the same as routine operations. The lymph nodes of group 5 and 8 on the surface of the common hepatic artery were replaced by proliferated vascular tissue. In case 1, left gastric artery and vein were not found, but notably increased artery blood supply on the upper edge of the lesser curvature was observed, which

could be the compensatory hypertrophic arteries after liver transplantation. Additionally, the patient suffered from fundus esophageal varices, which also increased the difficulty of lymph node dissection in groups 1, 3 and 7. The reconstruction of digestive tract was completed with the assistance of robot, which provided the flexibility by using robotic endoscopic joints.

In case 1, 52 lymph nodes were dissected, but only 19 were confirmed by pathologist, as the others were locally proliferated adipose tissue. In case 2, 44 lymph nodes were isolated, but only 21 were analyzed by pathologist after operation, and the number was much lower compared with primary GC. The reason could be reduction of lymph nodes in the abdominal, perihepatic, upper and lower pyloric lymph node groups after liver transplantation, which was replaced by locally proliferated adipose fibrous tissue.

Postoperative management was similar as primary GC. The effects of immunosuppressive agents on wound healing should be minimized. In order to prevent acute rejection without inhibiting the anti-tumor activity excessively, the concentration of immunosuppressive agents should be fixed during the operation to maintain normal immune activity (18). In case 2, immunosuppressive agents were continuously provided during the surgery at the original dosage.

Robot surgery system with flexible joints enables us to operate in narrow spaces, and it is broadly used in isolating the adhesions between liver and colon/duodenum and dissecting the adhesions formed by impaired common hepatic artery which loses its vascular sheath. This study provides novel insight on the development of GC surgery after liver transplantation using robotic system. However, we also need more cases and longer follow-up time to support the conclusion.

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Footnote

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Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at http://dx.doi.

org/10.21037/tcr-19-1786). The authors have no conflicts of interest to declare.

Ethical Statement: The author is 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. Written informed consent was obtained from the patient for publication of this manuscript and any accompanying images.

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Translational Cancer Research, Vol 9, No 6 June 2020

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