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ORIGINAL ARTICLE

Endoscopic ultrasound-guided hepaticogastrostomy for advanced cholangiocarcinoma after failed stenting by endoscopic retrograde cholangiopancreatography



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Received 14 November 2012; received in revised form 29 January 2013; accepted 12 April 2013
Available online 13 June 2013

KEYWORDS

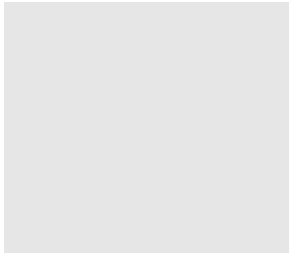
advanced
cholangiocarcinoma;
endoscopic
ultrasound;
hepaticogastrostomy

Summary *Objective:* Cholangiocarcinoma is common in Thailand. There are many palliative treatments available for patients with unresectable tumor, such as endoscopic retrograde cholangiopancreatography (ERCP) with stents, percutaneous transhepatic biliary drainage, or surgery. In cases in which ERCP has failed, we propose an alternative technique: the use of endoscopic ultrasound with fluoroscopy to perform hepaticogastrostomy for palliative drainage instead of percutaneous transhepatic biliary drainage.

Patients and methods: A case series study was conducted between December 2005 and December 2009 of 10 patients (4 male and 6 female, average age: 57 years) who presented with severe jaundice caused by advanced cholangiocarcinoma, who were treated with this procedure after failure to drain by ERCP. We used an electronic convex curved linear-array fluoroscopy-guided echoendoscope to drain the left dilated intrahepatic duct to the stomach by metallic wallstent. We performed the procedure with the first six patients under general anesthesia and with the other four under conscious sedation. Follow-up liver function tests were done, and clinical symptoms and survival times were recorded.

Results: Hepaticogastrostomy was unsuccessful on the first two patients (success rate = 8/10; 80%), and effective drainage was obtained in only seven patients. Average total bilirubin reduction was 14.96 mg/dL (58.75%) and 18.13 mg/dL (71.20%) after 2 weeks and 4 weeks, respectively, with good quality of life. One patient was not effectively drained because of malposition of the stent. There were two patients whose stent migrated into the stomach;

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one needed a second session with a second wallstent, and the other needed a double pigtail stent inside the second wallstent. Follow-up survival rates were 32–194 days (average: 123 days).

Conclusion: Endoscopic-ultrasound-guided hepaticogastrostomy is safe and can be a good palliative option for advanced malignant biliary obstruction because it drains internally and is remote from the tumor site, promoting a long patency period of prosthesis and better quality of life.

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1. Introduction

Cholangiocarcinoma is a common malignant tumor in the Thai population, especially in the northeastern part of Thailand; an area in which liver fluke is endemic. The curative treatment is surgery, but most patients are inoperable by the time they consult a specialist. The choice of treatment for patients with biliary obstruction caused by inoperable cancer is controversial. Usually, the first choice is endoscopic retrograde cholangiopancreatography (ERCP) with plastic or metallic stents. Another palliative treatment is percutaneous transhepatic biliary drainage (PTBD), including external drainage or internal drainage by radiologically placing a stent through the tumor into the common bile duct; however, ~28% of these patients have recurrent obstruction and cholangitis.¹ Palliative hepaticocentric surgical bypass may offer longer-term patency and better quality of life compared with PTBD, but the patients will suffer from postoperative pain, complications and higher mortality. This proposed technique instead of PTBD, is also a palliative procedure that drains the biliary duct far from the site of tumor and avoids open surgery by using an endoscopic ultrasound (EUS)-guided method to perform hepaticogastrostomy as a minimally invasive option.^{2–8}

2. Patients and methods

2.1. Patients

A case series study of minimally invasive trial EUS-guided hepaticogastrostomy was conducted from December 2005 to



Figure 1 Computed tomography scan of the liver showed marked dilated bilateral intrahepatic bile duct in type 4 hilar cholangiocarcinoma.

December 2009 on 10 patients with indications including obstructive jaundice from inoperable (advanced) hilar cholangiocarcinoma with failed ERCP and significant dilation of the left intrahepatic bile duct (Fig. 1). The exclusion criteria were shock, coagulopathy, ascites, and non-cooperation. Four of the patients were male and the other six were female, and their average age was 57 years. There were no pathologically proven cases of cholangiocarcinoma. All patients were diagnosed by clinical findings and imaging. We performed EUS-guided hepaticogastrostomy using the technique described below (Figs. 2 and 3); the first six patients received the procedure under general anesthesia and the other four under conscious sedation. The patients who drained successfully had follow-up progression of disease and blood level of liver function and alkaline phosphatase were followed 2 and 4 weeks after procedure and data of survival times were recorded.

2.2. Technique of EUS-guided hepaticogastrostomy

The EUS-guided hepaticogastrostomy technique introduced by M. Giovannini consists of a two-step procedure which creates an anastomosis between the dilated left intrahepatic bile duct and the stomach.² He placed a plastic stent in first step and then replaced it by a covered metallic stent a few weeks later. In this study, we used the Pentax EG 38 UT with EUB 6500 Hitachi Ultrasound with convex curved linear array echo-endoscope and C-arm fluoroscopy. This technique was applied from technique of EUS-FNA (fine needle aspiration) by using needle via the EUS endoscope to



Figure 2 Endoscopic ultrasound view of needle punctured through the gastric wall into the dilated left intrahepatic bile duct.

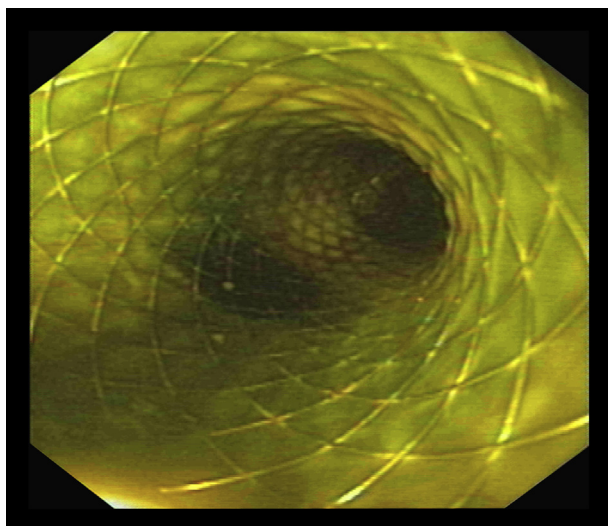


Figure 3 An 8 Fr 60-mm metallic wallstent in place of hepaticogastrostomy with bile flow.

create tract from lesser curve of stomach to left intrahepatic duct under general anesthesia or conscious sedation (25 mg pethidine + 2.5 mg midazolam), the EUS scope was introduced through the mouth into the stomach. Through the lesser curvature of the stomach, the dilated left intrahepatic duct in segment II was identified and punctured with a 19-gauge EUS needle (Wilson–Cook, Winston-Salem, NC, USA). Under fluoroscopic control, a contrast agent was injected to show the intrahepatic bile duct. A guide wire (0.035 inch diameter) was inserted into bile duct and a cystotome was applied via the scope to create a tunnel at the lesser curvature of the stomach wall, then a needle knife was applied via the scope to puncture

the liver into the bile duct by using cauterization. We preferred a single-step procedure, so instead of placing a plastic stent to create a fistula, after removing the needle knife, we placed a 10-mm diameter, 60-mm length covered, self-expanding metallic stent or wallstent (Boston Scientific Corporation, Boston, MA, USA) in one step and then removed the guide wire and EUS scope.

3. Results

EUS-guided hepaticogastrostomy was successful in eight of the 10 patients without any immediate complication such as massive pneumoperitoneum. Placement of stents was unsuccessful in the first two patients (success rate = 8/10; 80%). Effective drainage was obtained in seven patients while the stent position and drainage in the third patient was not so effective, because the penetration site of the stomach was at the esophagogastric junction. Follow-up observations after 2 weeks and 4 weeks found that average total bilirubin fell from 25.46 mg/dL to 10.50 mg/dL and 7.33 mg/dL (or reduction rate = 58.75% and 71.20%), respectively (Table 1). All successful patients had a good recovery, and started an oral diet on the day after the procedure. Follow-up computed tomography (CT) scans after 2 weeks showed metallic stents, with marked reduction of dilated intrahepatic bile duct (Fig. 4). There were two patients whose stents were dislodged and migrated into the stomach, detected by plain abdominal X-ray on the day after the procedure; one needed a second session with a second wallstent and the other required a second wallstent and a double pigtail stent placed inside the wallstent to prevent migration due to the tunnel being too large. Most of the followed up patients had much improved jaundice and a good quality of life, and the survival times were

Table 1 Patient data before and after intervention.

Patient	Sex	Age (y)	Hb (g/dL)	WBC/ μ L	CEA (ng/mL)	Bismuth type	Pre-intervention		2 wk post-intervention		4 wk post-intervention	
							Bilirubin (TB/DB) (mg/dL)	AP (μ L)	Bilirubin (TB/DB) (mg/dL)	AP (μ L)	Bilirubin (TB/DB) (mg/dL)	AP (μ L)
1	M	51	9.3	7500	3.15	3A	25.44/20.14	518	27.29/20.29	1269	27.46/18.78	1062
2	F	59	10.6	6800	6.22	4	20.50/17.48	488	3.42/2.16	358	1.25/0.64	615
3	F	44	10.2	10,900	5.38	4	38.37/29.45	494	10.42/7.87	501	5.02/3.69	1158
4	M	48	9.9	9300	7.90	4	22.00/16.09	521	12.77/8.80	489	5.20/2.50	720
5	F	62	11.5	11,000	3.06	3A	22.62/16.78	655	4.22/3.19	218	0.72/0.38	60
6	F	56	10.8	5700	12.18	4	17.60/9.10	823	18.88/13.32	200	14.55/7.10	253
7	M	67	12.6	6300	9.05	4	26.31/18.99	1427	10.07/5.92	716	10.62/7.37	1863
8	F	60	10.3	5500	1.96	4	21.83/16.54	484	18.00/14.50	255	8.55/4.92	220
9	M	58	11.7	8800	3.44	3A	66.27/40.46	2726	10.50/7.60		7.33/4.53	
10	F	65	9.2	17,200	5.38	4	39.64/27.49	443	14.96 (58.75%)		18.13 (71.2%)	
Average		57	10.6	7800	5.77		25.46/17.49					
Reduction of total bilirubin												

AP = alkaline phosphatase; CEA = carcinoembryonic antigen; DB = direct bilirubin; F = female; Hb = hemoglobin; M = male; TB = total bilirubin; WBC = white blood cell count.



Figure 4 Follow-up computed tomography scan showed metallic stent drained at lesser curvature of the stomach and nearly normal size of the bile duct.

32–194 days (average: 123 days) (Table 2). The causes of death were mostly advanced stage of the disease with liver or multiorgan failure.

4. Discussion

Cholangiocarcinoma is a common malignant tumor of the bile duct in the Thai population and has the highest recorded incidence in the world in Northeast Thailand. For resectable cases, the treatment of choice is surgical resection but most patients present when the disease is at an advanced stage or inoperable. There are several options for treatment, with ERCP and endoprosthesis being the first choice. If ERCP fails or is not possible, PTBD or endoprosthesis trans-PTBD are other options but PTBD has a high risk of infection, obstruction, and subsequent poor quality of life because the stent is placed across the tumor and the

catheter drains into a bag outside the body. Becker et al have demonstrated a higher rate of complications in patients with hilar involvement compared with those without hilar involvement, with significantly lower stent patency rates.¹

There are many techniques for draining the biliary duct far from the tumor site, as described by Soulez et al using fluoroscopic, endoscopic and laparoscopic guidance to create a hepaticogastric anastomosis,⁹ and Tipaldi using CT guidance and fluoroscopy with two steps of percutaneous hepaticogastric drainage.¹⁰ Warren indicated that diversion of bile from the liver into the stomach does not cause gastric ulcers, which is another advantage of hepaticogastrostomy,¹¹ but Moss has indicated that gastritis could be a potential complication due to the contact of gastric mucosa with bile.¹²

Endoscopic ultrasound, which was developed >20 years ago, is currently a valuable diagnostic instrument and also a therapeutic intervention. Since the introduction of the curved linear array echoendoscope in 1990, many procedures such as cystogastrostomy for pancreatic pseudocyst, choledochoduodenostomy and hepaticogastrostomy can be performed by minimally invasive techniques.^{2–8} EUS-guided hepaticogastrostomy has the same level of safety as compared with PTBD, but higher effectiveness of drainage because of the larger diameter of the metallic stent than the PTBD catheter (10 mm vs. 3 mm).

With our first two patients, in the early stages when we had no previous experience, we failed to place the stent because of technical error and we consulted with a radiologist to perform PTBD in both cases. With the third patient, we succeeded in placing the stent, but the position of the stent was not favorable because it drained from the left intrahepatic duct through the esophagogastric junction, so that it was at the same level as the bile duct and the bile did not drain well by the natural effect of gravity. However, the CT scan showed much improvement in the dilated bile duct after 2 weeks (Fig. 5). With the other patients, with the benefit of more experience, we placed the stents more effectively, and bilirubin levels fell rapidly. At the 2-week follow-up observation, there was significant improvement in the patients' bile ducts, and they had a good appetite and a better quality of life.

Table 2 Success rate, complications, survival time and cause of death.

Patient	Result of procedure	Complications	Survival time	Cause of death
1	Failure	—	—	—
2	Failure	—	—	—
3	Success	Cholangitis	1 m 2 d	Sepsis
4	Success	—	6 m 10 d	Sepsis
5	Success	Stent migration	7 m 24 d	Hepatic failure
6	Success	—	3 m 4 d	Hepatic failure
7	Success	—	4 m 10 d	Hepatic failure
8	Success	—	1 m 11 d	Hepatic failure
9	Success	Stent migration	5 m 2 d	Sepsis
10	Success	—	5 m 1 d	Hepatic failure
Average			123 d (4 mo)	



Figure 5 Follow-up computed tomography scan showed metallic stent drained at the esophagogastric junction and much smaller size of the bile duct in the third patient.

The complications and prognostic factors for mortality in these patients were advanced malignancy, impaired liver function, off and on cholangitis, bile leakage to the peritoneal cavity, stent obstruction or malposition, and poor drainage of bile. Two patients (Cases 3 and 8) that were not well drained had survival time of only ~1 month because they quickly developed sepsis and hepatic failure. In other patients, with rapid reduction of bilirubin, appropriate positioning of the stent, and very good drainage of bile, longer survival times were achieved. There were no definite episodes of bile leakage and stent obstruction.

In conclusion, EUS-guided hepaticogastrostomy can be done safely, with short hospitalization, and a longer period of stent patency. It makes the patients more comfortable and results in a better quality of life. This technique should be an alternative to PTBD or surgical drainage for patients with advanced cholangiocarcinoma when palliative ERCP fails or is not possible.

Acknowledgments

Assoc. Prof. Sukij Panpimanmas, accepts full responsibility for the conduct of this study, collection and/or interpretation of data, and/or drafting the manuscript. Dr Thawee Ratanachu-ek has approved the final draft submitted.

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