Intrahepatic cholangiocarcinoma diagnosed via endoscopic retrograde cholangiopancreatography with a short double-balloon enteroscope

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Abstract
Endoscopic retrograde cholangiopancreatography (ERCP) using a double-balloon enteroscope (DBE) in patients with bowel reconstruction due to a previous abdominal surgery is now widely accepted. In particular, a short DBE, which has a 2.8-mm working channel and 152-cm working length, is useful for ERCP because of its good rotational and straightening ability and the availability of various conventional ERCP accessories through the working channel. Herein we report a case of intrahepatic cholangiocarcinoma via ERCP with a short DBE. This is the first report in which the pre-cutting and the brush cytological examination were performed successfully under a DBE to diagnose intrahepatic cholangiocarcinoma pathologically. The short DBE allowed us to perform all diagnostic and therapeutic procedures accepted in conventional ERCP in patients with surgically altered anatomies.

Key words: Double-balloon enteroscope; Diagnosis; Endoscopic retrograde cholangiopancreatography; Intrahepatic cholangiocarcinoma; Cytology

Core tip: This is the first report in which the pathological diagnosis of intrahepatic cholangiocarcinoma could be made using the brush cytological examination via endoscopic retrograde cholangiopancreatography with a double-balloon enteroscope (DB-ERCP). In this paper, the methods of these procedures during DB-ERCP such as biliary cannulation, pre-cutting, and cytological examination are demonstrated in detail. Therefore, we believe that this paper must expand awareness of the utility of DB-ERCP for diagnosis of pancreatobiliary disease in patients with bowel reconstruction due to a previous abdominal surgery.

INTRODUCTION
Performing endoscopic retrograde cholangiopancreatography (ERCP) using a conventional duodenoscope in patients with bowel reconstruction due to a previous abdominal surgery is challenging because it is frequently
impossible to reach the papilla or the hepatico/choledochojejunul and pancreaticojejunal anastomosis owing to the insufficient scope length and unusual postsurgical conditions such as intestinal adhesions and anastomosis angulation. However, the use of a double-balloon enteroscope (DBE), which was originally developed for the management of small bowel diseases, has been shown to make it more feasible to perform ERCP in patients with bowel reconstruction. Recently, ERCP using a DBE (DB-ERCP) is now widely accepted. In particular, a short DBE, which has a 2.8-mm working channel and 152-cm working length, is useful for ERCP because of its good rotational and straightening ability and the availability of various conventional ERCP accessories through the working channel. ERCP using a short DBE was reported to be useful and safe for diagnostic and therapeutic interventions such as sphincterotomy, stone extraction, and stent placement. In this paper, we report a case of a post-surgical patient diagnosed as having intrahepatic cholangiocarcinoma via ERCP with a short DBE. We aimed to demonstrate the feasibility of successfully achieving access to the biliary duct by pre-cutting under a DBE and obtaining the pathological diagnosis by tissue sampling via DB-ERCP.

CASE REPORT

A 74-year-old man, who had undergone partial gastrectomy with gastrojejunostomy and Roux-en-Y reconstruction for gastric carcinoma five years before, presented to another hospital to evaluate a mass lesion in hepatic segment VII. A fluorine-18-fluorodeoxyglucose positron emission tomography scan revealed a pathological uptake pattern with a maximum standardized uptake value of 5.6 in the mass, suggestive of a malignant tumor. An ultrasound-guided percutaneous needle biopsy was performed to obtain a pathological diagnosis of the mass. However, the biopsy finding revealed no sign of malignancy. In addition, the patient developed biliary peritonitis after the liver biopsy and therefore required hospitalization and conservative treatment for approximately 1 mo. One month after discharge from the hospital, an ultrasound image revealed the mass lesion remained constant in size; however biliary stricture with upstream dilatation, which appeared to result from the mass lesion, was noted on MRCP (Figure 1A). On MRCP, a focal biliary stricture with upstream dilatation in the branch arising from the right anterior duct was observed (Figure 1B). The patient underwent ERCP with a short DBE (EC-450B15; Fujifilm, Osaka, Japan) in the prone position with carbon dioxide (CO₂) insufflations. At first, a short DBE was carefully inserted into the blind loop and reached the papilla. Subsequently, standard biliary cannulation using a straight cannula (PR-10Q; Olympus, Tokyo, Japan) was attempted after the papilla was moved to the lower endoscopic field of view by keeping the overtube balloon inflated and rotating the enteroscope (Figure 2A), as described in a previous report. However, we were unable to obtain cholangiography, but only pancreatography. Therefore, a 0.025-inch guide wire (Jagwire; Boston Scientific Japan, Tokyo, Japan) was inserted into the main pancreatic duct (Figure 3A), and the pre-cut technique was carried out with a wire-guided sphincterotome (Autotome RX, Boston Scientific, Japan). The incision was started toward the biliary direction and was stopped at the lower one-third of the ampullary mound (Figure 2B). After the incision, the orifice of the biliary duct became visible (Figure 2C), and selective biliary cannulation with a straight cannula was performed successfully (Figure 3B). After contrast enhancement of the biliary duct, a focal stricture in the branch of the right intrahepatic duct was identified under radiographic guidance (Figure 3C). A guide wire followed by a cannula was successfully passed through the stricture after several negotiations (Figure 3D). Subsequently, tissue sampling for cytological examination was performed at the level of the stricture using a brush device (Cytomax II DLB 35-1.5; Cook, Osaka, Japan) introduced by a guide wire (Figure 3E). Finally, the nasobiliary drainage tube was placed at the distal side to collect the bile for cytological examination (Figure 3F). The day after the ERCP, a slight increase in the serum amylase level was observed, but the patient did not complain of abdominal pain. The result of the cytological examinations of the sample obtained by the brush and nasobiliary drainage tube was positive for malignancy. On the basis of the cytological finding, the patient underwent segmentectomy of the hepatic cholangiocarcinoma.
left lobe. The final pathological report indicated intrahepatic cholangiocarcinoma.

**DISCUSSION**

With the advent of DBEs, the endoscopic management of pancreaticobiliary diseases has become more feasible in patients with previous gastrointestinal surgery such as Billroth II gastrojejunostomy, Roux-en-Y reconstruction, and Whipple’s resection\(^8\)\(^{-6}\). Accordingly, in these years, the number of published reports on the utility of DB-ERCP has been increasing. In most of these previous reports, a standard DBE (EN-450T5; Fujifilm)\(^8\)\(^{-6}\) and a single-balloon enteroscope\(^7\) were used. However these enteroscopes have a limitation with regard to the availability of its accessories because of its 200-cm working length. In contrast, a short DBE has a 2.8-mm working channel and 152-cm working length, for which various diagnostic and therapeutic accessories used in conventional ERCP, including sphincterotome, balloon catheter, basket, biopsy forceps, brush, intraductal ultrasonic probe and biliary stent, are available without modification\(^8\)\(^{-11}\).
The use of a short DBE appears to not only widen the parameters of the procedures which we can perform in DB-ERCP, but also increase the success rate of the procedure owing to its good maneuverability. In fact, with a short DBE, we reported higher success rates in deep insertion [100/103 (97%)], cholangiography [98/100 (98%)], and therapeutic interventions [98/98 (100%)], compared with the previous reports.

Deep cannulation to the biliary duct is the most important step for successful ERCP. The failure rate for cannulation of the naive papilla in conventional ERCP is reported to be up to 10% of the ERCP procedures.[12,13] In general, this rate in DB-ERCP might be even higher because of the difficult location of the papilla and the instability of the manipulation of the enteroscope and cannula. In the case of a difficult biliary cannulation in conventional ERCP, the pre-cut techniques have been used as a widely accepted option to facilitate the biliary access.[14,15] The pre-cut techniques include needle knife sphincterotomy and wire assisted transpancreatic sphincterotomy. In contrast with conventional duodenoscopy, a DBE is forward viewing and has no elevator function, making the pre-cutting technically more challenging. From our experience, we recommend wire-guided transpancreatic sphincterotomy in performing the pre-cutting under a DBE, because the direction, depth, and length of incision are easier to control, compared with needle knife papillotomy performed freehand.

Although intrahepatic stricture frequently results from malignant disease such as cholangiocarcinoma and hepatocellular carcinoma, it is important to confirm the malignant cells histopathologically by the tissue sampling method, because an accurate diagnosis is crucial for the choice of an appropriate therapeutic strategy. In general, percutaneous needle biopsy is accepted for histological diagnosis when the mass causing the intrahepatic stricture can be detected on ultrasonography. However, the potential disadvantage of this technique is that complications such as pneumothorax, hemorrhage, biliary leakage, biliary peritonitis, and tumor seeding can occur.[6-17] In contrast, the transpapillary tissue sampling methods during ERCP include brush cytological examination, bile cytological examination, and biopsy. In particular, brush cytological examination is commonly used because it is relatively simple and safe. Although the specificity of this method for the diagnosis of cholangiocarcinoma is almost 100%, the sensitivity is by no means satisfactory, ranging from 44% to 81%.[18,19] In our institution, when brushing is performed, bile cytological examination is also performed via a nasobiliary drainage tube placed subsequently after brushing. We believe that such a combination of specimen collection methods enhances the cancer detection rate.

In conclusion, the short DBE allowed us to perform all diagnostic and therapeutic procedures accepted in conventional ERCP in patients with surgically altered anatomy.

REFERENCES

1 Yamamoto H, Sekine Y, Sato Y, Higashizawa T, Miyata T, lino S, Ido K, Sugano K. Total endoscopy with a nonsurgically steerable double-balloon method. Gastrointest Endosc 2001; 53: 216-220 [PMID: 11174299]

2 Charpentier B, Hiesse C, Pitran C, Lantz O, Fries D, Bach JF, Chatenoud L. Acute clinical syndrome associated with OKT3 administration. Prevention by single injection of an anti-human TNF monoclonal antibody. Presse Med 1991; 20: 2009-2011 [PMID: 1837113 DOI: 10.1111/j.1572-0241.2007.01745.x]

3 Chu YC, Yang CC, Yeh YH, Chen CH, Yueh SK. Double-balloon enteroscopy application in biliary tract disease-its therapeutic and diagnostic functions. Gastrointest Endosc 2008; 68: 855-591 [PMID: 18561917 DOI: 10.1016/j.gie.2008.03.1083]

4 Aabakken L, Breithauer M, Line PD. Double-balloon enteroscopy for endoscopic retrograde cholangiography in patients with a Roux-en-Y anastomosis. Endoscopy 2007; 39: 1068-1071 [PMID: 18072058]

5 Mönkemüller K, Fry LC, Bellutti M, Neumann H, Malfertheiner P. ERCP with the double balloon enteroscope in patients with Roux-en-Y anastomosis. Surg Endosc 2009; 23: 1961-1967 [PMID: 1967052 DOI: 10.1007/s00464-008-0229-8]

6 Raithel M, Dormann H, Naegel A, Boxberger F, Hahn EG, Neurath MF, Maiss J. Double-balloon-enteroscopy-based endoscopic retrograde cholangiopancreatography in post-surgical patients. World J Gastroenterol 2011; 17: 2302-2314 [PMID: 21633596 DOI: 10.3748/wjg.v17.i18.2302]

7 Wang AY, Sauer BG, Behm BW, Ramanath M, Cox DG, Ellen KL, Shami VM, Kahaleh M. Single-balloon enteroscopy effectively enables diagnostic and therapeutic retrograde cholangiography in patients with surgically altered anatomy. Gastrointest Endosc 2010; 71: 641-649 [PMID: 20189529 DOI: 10.1016/j.gie.2009.10.051]

8 Shimatani M, Matsuhashita M, Takaoka M, Koyabu M, Ikeura T, Kato K, Fukui T, Uchida K, Okazaki K. Effective “short” double-balloon enteroscope for diagnostic and therapeutic ERCP in patients with altered gastrointestinal anatomy: a large case series. Endoscopy 2009; 41: 849-854 [PMID: 19750447 DOI: 10.1055/s-0029-1215108]

9 Shimatani M, Matsuhashita M, Takaoka M, Kusuda T, Fukuta N, Koyabu M, Uchida K, Okazaki K. “Short” double-balloon enteroscope for endoscopic retrograde cholangiopancreatography with conventional sphincterotomy and metallic stent placement after Billroth II gastrectomy. Endoscopy 2009; 41 Suppl 2: E19-E20 [PMID: 19219763 DOI: 10.1055/s-0028-1103466]

10 Matsuhashita M, Shimatani M, Takaoka M, Okazaki K. “Short” double-balloon enteroscope for diagnostic and therapeutic ERCP in patients with altered gastrointestinal anatomy. Am J Gastroenterol 2008; 103: 3218-3219 [PMID: 19086985 DOI: 10.1111/j.1572-0241.2008.02161_18.x]

11 Itoi T, Ishii K, Sofuni A, Itokawa F, Tsuchiya T, Kurihara T, Tsuji S, Ikeuchi N, Futakawa K, Moriyasu F, Tsuchida A. Long- and short-type double-balloon enteroscopy-assisted therapeutic ERCP for intact papilla in patients with a Roux-en-Y anastomosis. Surg Endosc 2011; 25: 713-721 [PMID: 20976503 DOI: 10.1007/s00464-010-1226-4]

12 Cortas GA, Mehta SN, Abraham NS, Barkun AN. Selective cannulation of the common bile duct: a prospective randomized trial comparing standard catheters with sphincterotomes. Gastrointest Endosc 1999; 50: 775-779 [PMID: 10570335]

13 Schwacha H, Allgaier HP, Deibert P, Olschewski M, Allgaier U, Blum HE. A sphincterotome-based technique for selective transpapillary common bile duct cannulation. Gastrointest Endosc 2000; 52: 387-391 [PMID: 10968855]

14 Testoni PA, Testoni S, Giussani A. Difficult biliary cannulation during ERCP: how to facilitate biliary access and minimize the risk of post-ERCP pancreatitis. Dig Liver Dis 2011; 43: 596-603 [PMID: 21377432 DOI: 10.1016/j.dld.2011.01.019]

15 Wang P, Zhang W, Liu F, Li ZS, Ren X, Fan ZN, Zhang X,
Lu NH, Sun WS, Shi RH, Li YQ, Zhao Q. Success and complication rates of two precut techniques, transpancreatic sphincterotomy and needle-knife sphincterotomy for bile duct cannulation. *J Gastrointest Surg* 2010; 14: 697-704 [PMID: 20054659 DOI: 10.1007/s11605-009-1134-x]

Piccinino F, Sagnelli E, Pasquale G, Giusti G. Complications following percutaneous liver biopsy. A multicentre retrospective study on 68,276 biopsies. *J Hepatol* 1986; 2: 165-173 [PMID: 3958472]

Chang S, Kim SH, Lim HK, Lee WJ, Choi D, Lim JH. Needle tract implantation after sonographically guided percutaneous biopsy of hepatocellular carcinoma: evaluation of doubling time, frequency, and features on CT. *AJR Am J Roentgenol* 2005; 185: 400-405 [PMID: 16037512]

de Bellis M, Sherman S, Fogel EL, Cramer H, Chappo J, McHenry L, Watkins JL, Lehman GA. Tissue sampling at ERCP in suspected malignant biliary strictures (Part 2). *Gastrointest Endosc* 2002; 56: 720-730 [PMID: 12397282]
