Endoscopic retrograde cholangiopancreatography in periampullary diverticulum: The challenge of cannulation

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Abstract
Periampullary diverticulum (PAD) is duodenal outpunching defined as herniation of the mucosa or submucosa that occurs via a defect in the muscle layer within an area of 2 to 3 cm around the papilla. Although PAD is usually asymptomatic and discovered incidentally during endoscopic retrograde cholangiopancreatography (ERCP), it is associated with different pathological conditions such as common bile duct obstruction, pancreatitis, perforation, bleeding, and rarely carcinoma. ERCP has a low rate of success in patients with PAD, suggesting that this condition may complicate the technical application of the ERCP procedure. Moreover, cannulation of PAD can be challenging, time consuming, and require the higher level of skill of more experienced endoscopists. A large portion of the failures of cannulation in patients with PAD can be attributed to inability of the endoscopist to detect the papilla. In cases where the papilla is identified but does not point in a suitable direction for cannulation, different techniques have been described. Endoscopists must be aware of papilla identification in the presence of PAD and of different available cannulation techniques, as well as the technical feasibility and safety of each.

Key words: Periampullary diverticulum; Cannulation techniques; Tips; Endoscopic ultrasound; Endoscopic retrograde cholangiopancreatography

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INTRODUCTION

Periampullary diverticulum (PAD) is duodenal outpunching defined as herniation of the mucosa or submucosa that occurs via a defect in the muscle layer within an area of 2 to 3 cm around the papilla. Prevalence of PAD increases with age, and overall prevalence among the elderly is reportedly 65%. The formation of PAD is related to progression of duodenal motility disorders. Furthermore, increased intraduodenal pressure and progressive weakening of intestinal smooth muscles are known as the main underlying etiologies for this defect. PAD is sub-classified into two categories according to the location of the papilla with respect to the diverticulum. In type I, or peri-diverticular papilla, the papilla is located at the edge of the diverticulum or within a radius of 2 cm from the diverticular edge. In type II, or intradiverticular papilla (IDP), the papilla is located inside the diverticulum or lying between two adjacent diverticula.

Although PAD is usually asymptomatic and discovered incidentally in patients during endoscopic retrograde cholangiopancreatography (ERCP), it is associated with different pathological conditions such as common bile duct (CBD) obstruction, pancreatitis, perforation, bleeding, and rarely carcinoma. Several hypotheses have been put forth to explain the observed higher incidence of biliary stone formation in the presence of PAD. First, it was proposed that dysfunction in the sphincter of Oddi, which in turn causes reflux of pancreatic fluid and intestinal content, can lead to biliary stone formation. Second, it was proposed that diverticula cause spasm of the sphincter, thereby increasing biliary tract pressure that may in turn produce jaundice and cholangitis as well as predispose for cholecodolithiasis. Finally, it was proposed that PAD may compress the distal part of the CBD to cause functional biliary stasis, and this hypothesis was supported by the observation of increased incidence of pigment biliary stones.

Reported success rates of cannulation in patients with PAD have varied from 61% to 95.4%, a range that is significantly lower than that observed in patients without PAD. In recent years, new techniques and new devices for successful biliary cannulation have been developed to improve rates of success in patients with PAD. For patients with PAD, endoscopists must be aware of papilla identification and the different cannulation techniques available, including the technical feasibility and safety of each, in order to make an informed decision and ensure the best outcome. Herein, we review the literature on this practical topic that was obtained through an electronic search of the literature databases of Google Scholar and PubMed using the following terms alone or in combination: ERCP, difficult cannulation, cannulation techniques, and periampullary diverticulum.

TIPS FOR PAPILLARY ORIENTATION AND CANNULATION

The presence of PAD is thought to complicate the application of ERCP, an already technically difficult procedure. Cannulation of IDP can be challenging, time consuming and require the higher level of skill of more experienced endoscopists. A large portion of the failures of cannulation in patients with PAD has been attributed to inability of the endoscopist to detect the papilla. However, in some studies, the finding of PAD during an ERCP was suggested as an indicator of an easier cannulation attempt, with a reported success rate of 94.9% compared to that of 94.8% in non-PAD patients. After exclusion of cases with undetectable papillas that were considered to be likely IDPs, in ERCP, identification of the papilla is the first major obstacle, especially in the presence of large diverticula. Thus, it is extremely helpful to know the following tips: (1) in most cases, the papilla is located on the lower edge of the diverticulum or just inside, somewhere between the positions of 4 o'clock and 8 o'clock; (2) large diverticula are usually divided from proximal to distal by a ridge-like septum. This mostly involves the bile duct, with the ridge terminating at the papilla; (3) a catheter can be used to straighten and evert the folds to identify a hidden papilla within the diverticulum; (4) cannulation with the tip of the duodenoscope within the sac is also possible, but care must be taken to avoid perforation; and (5) in contrast to the usual papillary anatomy, the presence of PAD alters the biliary direction. It is often not acutely angulated superiorly, but runs more directly. Thus, acute angulation of the sphincterotome is not necessary.

TECHNIQUES FOR DIFFICULT CANNULATION

To address cases where the papilla is identified but does not point in a suitable direction for cannulation, the below-described techniques are available for consideration (Table 1).

Two-devices in one-channel method

A biopsy forceps is used to pull the duodenal mucosa adjacent to the papilla, bringing the papillary orifice out of the diverticulum. Another instrument, either a cannula or sphincterotome, is then inserted into the working channel of the endoscope together with the biopsy forceps. With coordination of the two instruments, biliary cannulation can be attempted (Figure 1A). A report of this technique applied to two PAD cases showed successful cannulation for both and with no complications in either (success rate...
Reversed guidewire method
A second guidewire is advanced in reverse (stiff end forward) through the working channel of the duodenoscope, alongside the sphincterotome. This wire is then used to push the mucosa adjacent to the papilla toward the lumen of the duodenum and to straighten the folds, anchoring the papilla in a better configuration and creating a suitable direction for cannulation. A report of this technique applied to one PAD case showed successful cannulation with no complication (success rate 100%)\(^\text{[15]}\).

Double endoscope method
A forward-viewing gastroscope is inserted inside the diverticulum for better visualization of the papilla. A foreign body forceps is used to grasp the tissue just beside the papilla in order to bring it into a better orientation. The gastroscope holding the papilla is left in place, to avoid backsliding after opening of the forceps. A side-viewing duodenoscope is inserted alongside the gastroscope. With both endoscopes positioned simu-

### Table 1 Techniques for difficult cannulation

| Two-devices in one-channel method | Reversed guidewire method | Double endoscope method | Balloon dilation of the narrow diverticular neck | Endoclip-assisted cannulation | Cap-assisted cannulation | Pancreatic duct stent placement followed by pre-cut biliary sphincterotomy | Percutaneous ultrasound-guided rendezvous technique | EUS-guided rendezvous technique |

EUS: Endoscopic ultrasound.

100%\(^\text{[14]}\).

**Reversed guidewire method**
A second guidewire is advanced in reverse (stiff end forward) through the working channel of the duo-

**Figure 1 Techniques for difficult cannulation.** A: Two-devices in one-channel method; B: Double endoscope method; C: Balloon dilation of the narrow diverticular neck; D: Endoclip-assisted cannulation; E: Cap-assisted cannulation; F: Pancreatic duct stent placement followed by pre-cut biliary sphincterotomy; G: Percutaneous ultrasound-guided rendezvous technique; H: Endoscopic ultrasound-guided rendezvous technique.
Itaneously in the duodenum, the CBD can be cannulated (Figure 1B). A report of this technique applied to one PAD case showed successful cannulation with no complication (success rate 100%)\(^\text{(16)}\).

**Balloon dilation of the narrow diverticular neck**
In narrow-necked papillary diverticula with the papilla located in the fundus of the diverticulum, endoscopic balloon dilation of the narrow diverticular neck, using a 15-mm stone retrieval balloon, can be done safely, bringing the papillary orifice into view. Cannulation of the bile duct can be attempted without any complications (Figure 1C). A report of this technique applied to three PAD cases showed successful cannulation and no complications (success rate 100%)\(^\text{(17)}\).

**Endoclip-assisted cannulation**
One or more endoclips can be used to rotate the IDP externally and to fix it on the outside rim of the diverticulum. This manipulation can successfully evert and fix the papilla on the diverticular margin in a better position, resulting in successful biliary cannulation (Figure 1D). A report of this technique applied to two PAD cases showed successful cannulation with no complications (success rate 100%)\(^\text{(18)}\).

**Cap-assisted cannulation**
A transparent cap is attached to the tip of a forward-viewing endoscope. At first, selective biliary cannulation can be attempted through the papillary orifice. If selective biliary cannulation fails, endoscopic fistulotomy can be attempted. Fistulotomy is performed between the lower two-thirds and the upper one-third of the papillary roof. To gain biliary access after the fistulotomy, the lower two-thirds and the upper one-third of the diverticulum. This manipulation can successfully evert and fix it on the outside rim of the diverticulum. One or more endoclips can be used to rotate the IDP externally and to fix it on the outside rim of the diverticulum. This manipulation can successfully evert and fix the papilla on the diverticular margin in a better position, resulting in successful biliary cannulation (Figure 1D). A report of this technique applied to two PAD cases showed successful cannulation with no complications (success rate 100%)\(^\text{(17)}\). A report of this technique applied to two PAD cases showed successful cannulation with no complications (success rate 100%)\(^\text{(17)}\).

**Pancreatic duct stent placement followed by pre-cut biliary sphincterotomy**
In the case of pancreatic duct cannulation, placement of a main pancreatic duct stent keeps the papilla out of the diverticulum, thereby facilitating pre-cut needle knife sphincterotomy and selective cannulation of the CBD (Figure 1F). A report of this technique applied to eight cases showed successful cannulation in seven of the patients (success rate 87.5%), with two of those requiring a second ERCP for success. In addition, two patients developed post-ERCP pancreatitis (complication rate 25%)\(^\text{(20)}\).

**Percutaneous ultrasound-guided rendezvous technique**
After the percutaneous ultrasound-guided transhepatic biliary puncture is performed a sterile guidewire is inserted into the CBD, then into the papilla. A snare or forces is then used to grasp the guidewire and pull it back through the working channel of the duodenoscope for subsequent over-the-wire cannulation (Figure 1G)\(^\text{(21)}\). However, it is sometimes difficult to grasp the guidewire, which may be damaged or kinked, during the withdrawal through the working channel of the duodenoscope; thus, passing a catheter over it is difficult or sometimes impossible\(^\text{(22)}\). A study on the percutaneous-ultrasound guided rendezvous technique applied to a total of fourteen patients showed success in 13 (success rate 93%) with complication (retroperitoneal perforation) experienced in only 1 (complication rate 7%)\(^\text{(21)}\).

**Endoscopic ultrasound-guided rendezvous technique**
When the echoendoscope is positioned in the stomach or duodenum, and the bile ducts can be visualized by the endoscopic ultrasound (EUS), a 19-gauge or 22-gauge needle are used to puncture the bile ducts. After aspiration of bile, contrast is injected through the EUS needle to facilitate display the intra- and extra-hepatic bile ducts. After confirmation of bile duct puncture, a guidewire is advanced distally through the CBD and across the papilla under fluoroscopic guidance. The endoscope exchange is performed after passage of the guidewire through the papilla into the duodenum. In this process, the echoendoscope is removed, leaving the guidewire in place, after which a duodenoscope is passed up to the papilla alongside the EUS-placed guidewire. Finally, a snare or forceps is used to grasp the guidewire and pull it back out of the working channel of the duodenoscope for subsequent over-the-wire cannulation. After access to the CBD is achieved, a standard ERCP can be performed (Figure 1H). A study on the EUS-guided rendezvous technique applied to a total of 45 patients showed success in 36 (success rate 80%) with complications (bile leakage and pneumoperitoneum) experienced in only 2 (complication rate 4%)\(^\text{(23)}\).

**PROPOSED ALGORITHM**
We propose an algorithm based on the previous techniques to increase the success rate of cannulation (Figure 2). It is important to note, however, that this algorithm has several limitations. First, it is based on a small number of published cases for most of the techniques. Second, the success rates are comparable in most of the techniques and the choice depends on the endoscopist’s preference and experience. Finally, percutaneous ultrasound-guided and EUS-guided rendezvous techniques are not available in all centers.

**Feasibility and safety of therapeutic maneuvers**
When therapeutic maneuvers are performed in patients with PAD the potential risks of complications are a concern, primarily because of the thin mucosa and the absence of sphincter muscle present in the ampullary area\(^\text{(24)}\). Currently, endoscopic papillary large balloon dilation (EPLBD) combined with limited endoscopic...
sphincterotomy (ES) (EPLBD + ES) is regarded as an effective maneuver for treating difficult CBD stones. It has been reported that perforation and hemorrhage are less frequent in cases treated with EPLBD + ES than in those treated with standard ES alone\(^{25,26}\). The tendency toward a shorter ballooning time in patients with PAD can be explained by the lack of sphincter muscle and the ease of ampullary widening facilitated by EPLBD, which suggest that EPLBD is a safe method for retrieval of CBD stones in patients with PAD\(^{24}\). Moreover, the complication rates of ERCP are similar in patients with or without PAD and the therapeutic outcome is not affected by the presence of PAD\(^{3,7}\).

**CONCLUSION**

PAD represents a technical barrier to the successful application of ERCP. Cannulation of IDP can be challenging, time consuming and require the skill of more experienced endoscopists. In cases where the papilla is identified but does not point in a suitable direction for cannulation, a number of feasible techniques are available for consideration. Moreover, complication rates of ERCP are similar in patients with and without PAD, and therapeutic outcome is not affected by the presence of PAD.

**REFERENCES**

1. **Shemesh E**, Klein E, Czerniak A, Coret A, Bat L. Endoscopic sphincterotomy in patients with gallbladder in situ: the influence of periampullary duodenal diverticula. *Surgery* 1990; **107**: 163-166 [PMID: 20997455]

2. **Lobo DN**, Balfour TW, Ifikhar SY. Periampullary diverticula: consequences of failed ERCP. *Ann R Coll Surg Engl* 1998; **80**: 326-331 [PMID: 9849331]

3. **Boix J**, Lorenzo-Zúñiga V, Añaños F, Domènech E, Morillas RM, Gassull MA. Impact of periampullary duodenal diverticula at endoscopic retrograde cholangiopancreatography: a proposed classification of periampullary duodenal diverticula. *Surg Laparosc Endosc Percutan Tech* 2006; **16**: 208-211 [PMID: 16921297 DOI: 10.1097/0128689-200608000-00002]

4. **Oddo F**, Chevallier P, Souci J, Baque J, Buckley MJ, Fabiani P, Daine B, Cousenett A. [Radiologic aspects of the complications of duodenal diverticula]. *J Radiol* 1999; **80**: 134-140 [PMID: 10209709]

5. **Yoneyama F**, Miyata K, Ohta H, Takeuchi E, Yamada T, Kobayashi Y. Excision of a juxtapapillary duodenal diverticulum causing biliary obstruction: report of three cases. *J Hepatobiliary Pancreat Surg* 2004; **11**: 69-72 [PMID: 15754050 DOI: 10.1007/s00534-003-0854-7]

6. **Tyagi P**, Sharma P, Sharma BC, Puri AS. Periampullary diverticulitis and technical success of endoscopic retrograde cholangiopancreatography. *Surg Endosc* 2009; **23**: 1342-1345 [PMID: 18818967 DOI: 10.1007/s00464-008-0167-7]

7. **Panteris V**, Vezakis A, Filippou G, Filippou D, Karamanolis D, Rizos S. Influence of juxtapapillary diverticula on the success or difficulty of cannulation and complication rate. *Gastrointest Endosc* 2008; **68**: 903-910 [PMID: 1865174 DOI: 10.1016/j.gie.2008.03.1092]

8. **Yıldırım M**, Başoğlu M, Yılmaz O, Takeuchi E, Yamada T, Kobayashi Y. Excision of a juxtapapillary duodenal diverticulum causing biliary obstruction: report of three cases. *J Hepatobiliary Pancreat Surg* 2004; **11**: 69-72 [PMID: 15754050 DOI: 10.1007/s00534-003-0854-7]

9. **Hagège H**, Berson A, Pelletier G, Fritsch J, Choury A, Lizoury C, Etienne JP. Association of juxtapapillary diverticula with cholecloithiasis but not with cholecystolithiasis. *Endoscopy* 1992; **24**: 248-251 [PMID: 1612038 DOI: 10.1055/s-2007-101476]

10. **Miyazaki S**, Sakamoto T, Miyata M, Yamasaki Y, Yamashiki H, Kawata K. Function of the sphincter of Oddi in patients with juxtapapillary duodenal diverticula: evaluation by intraoperative bile manometry under a duodenal pressure load. *World J Surg* 1995; **19**: 307-312 [PMID: 7754640 DOI: 10.1007/BF00308647]

11. **Shinagawa N**, Fukui T, Mashita K, Kitano Y, Yura J. The relationship between juxtapapillary duodenal diverticula and the presence of bacteria in the bile. *Jpn J Surg* 1991; **21**: 284-291 [PMID: 1906956 DOI: 10.1007/BF00247940]

12. **Zoepf T**, Zoepf DS, Arnold JC, Benz C, Riemann JF. The relationship between juxtapapillary duodenal diverticula and disorders of the biliopancreatic system: analysis of 350 patients.
13. Pohl J. Periampullary Diverticulum: Cannulation and Sphincterotomy. Video J Encyclop GI Endosc 2013; 1: 516-517 [DOI: 10.1016/S2212-0971(13)70226-7]

14. Fujita N, Noda Y, Kobayashi G, Kimura K, Yaga A. ERCP for intradiverticular papilla: two-devices-in-one-channel method. Endoscopic Retrograde Cholangiopancreatography. Gastrointest Endosc 1998; 48: 517-520 [PMID: 9831843 DOI: 10.1016/S0016-5107(98)70096-3]

15. Elmunzer BJ, Boetticher NC. Reverse guidewire anchoring of the papilla for difficult cannulation due to a periampullary diverticulum. Gastrointest Endosc 2015; 82: 957 [PMID: 26142553 DOI: 10.1016/j.gie.2015.05.054]

16. Külling D, Haskell E. Double endoscope method to access intradiverticular papilla. Gastrointest Endosc 2005; 62: 811-812 [PMID: 16246708 DOI: 10.1016/j.gie.2005.06.035]

17. Tóth E, Lindström E, Fork FT. An alternative approach to the inaccessible intradiverticular papilla. Endoscopy 1999; 31: 554-556 [PMID: 10533741 DOI: 10.1055/s-1999-959]

18. Huang CH, Tsou YK, Lin CH, Tang JH. Endoscopic retrograde cholangiopancreatography (ERCP) for intradiverticular papilla: endoclip-assisted biliary cannulation. Endoscopy 2010; 42 Suppl 2: E223-E224 [PMID: 20931451 DOI: 10.1055/s-0029-1215008]

19. Myung DS, Park CH, Koh HR, Lim SU, Jun CH, Ki HS, Park SY, Rew JS. Cap-assisted ERCP in patients with difficult cannulation due to periampullary diverticulum. Endoscopy 2014; 46: 352-355 [PMID: 24549783]

20. Fogel EL, Sherman S, Lehman GA. Increased selective biliary cannulation rates in the setting of periampullary diverticula: main pancreatic duct stent placement followed by pre-cut biliary sphincterotomy. Gastrointest Endosc 1998; 47: 396-400 [PMID: 9609434 DOI: 10.1016/S0016-5107(98)70226-3]
