

How to perform EUS-based cholecystolithotomy

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ABSTRACT

EUS-guided gallbladder drainage (EUS-GBD) is gradually emerging as an option for patients with cholecystitis. A stent-bridged endoscopic intervention in the gallbladder (GB) has been proposed. The formation of a fistulous tract after EUS-GBD facilitates the entry of the endoscope inside the GB to perform endoscopic lithotomy and polypectomy, which is an efficient and safe procedure. However, the technical aspects of this procedure in order to optimize the success rate have only been scarcely discussed. This paper discusses the EUS-GBD prepared for per-oral cholecystolithotomy, the stent indwelling time, the procedure, the devices of per-oral cholecystolithotomy and the follow-up. The recently published literature on this topic is also updated in this review.

Key words: Cholecystolithotomy, EUS, gallbladder drainage

INTRODUCTION

EUS-guided gallbladder drainage (EUS-GBD) is a promising and novel method to treat acute cholecystitis in high-risk surgical patients.^[1-4] The technical and clinical success rates for EUS-GBD are reported to be more than 95%, with minor adverse events in <10% of the cases.^[5,6] However, after EUS-GBD, most of the patients are followed with a "wait-and-see" strategy until clinical symptoms or stent-related adverse events develop.^[7] For gallstones and gallbladder (GB) polys, further steps can be employed based on the drainage technique.

The path created by the stent can also serve as a conduit for a subsequent gallstone removal procedure.^[8,9] This minimally invasive technique can preserve the GBs that still have an adequate function. Furthermore, this

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GB-preserving cholecystolithotomy method facilitated by EUS and the metal stents might have more advantages in comparison to previous percutaneous or laparoscopic GB-preserving cholecystolithotomy methods.^[10,11]

However, the technical aspects of the procedure have been scarcely discussed. This paper the technical aspects of EUS-based cholecystolithotomy while providing an update of the recently published literature on the topic.

METHODS

This review first provides a brief summary of endoscopic cholecystolithotomy technique, which is based on the EUS-GBD. Then, this review is focused on the techniques for endoscopic cholecystolithotomy, which is

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presently very limited in the literature. We have searched PubMed using the following singular and combined search terms: <endoscopic ultrasound>; <gallbladder>; <drainage>; <gallstones>; <cholecystoscopy>; <Cholecystolithotomy>; <lumen-apposing metal stents>; <polyp-resection>; <cholecystitis>. Also, we have provided a discussion based on the experience from the tertiary endoscopic center.

RESULTS

The EUS-guided gallbladder drainage technique

The EUS-GBD technique started in 2007 when the double pigtail plastic stents or nasocystic drainage tube was used for the drainage. The EUS-GBD technique has undergone continuous improvement through the use of plastic stents, self-expandable metal stents, and a lumen apposing metal stent (LAMS); these continuous improvements have increased the success rate.^[12] The LAMSs can keep the target object closely opposed to the gastrointestinal (GI) wall. However, the stenting procedure is technically challenging and time consuming. Then, the LAMS with an electro-cautery enhanced delivery system (ECE-LAMS) was introduced and it significantly shortened the operation time.

The studies that refer to the learning curve of EUS-GBD indicate that endoscopists that have performed EUS-GBD are expected to achieve a reduction in the procedure time over successive cases. In terms of the efficiency, the procedure requires 41 min with a learning rate of 19 cases.^[13] However, the results were drawn from highly experienced and specialized endoscopists in the tertiary centers. For those who have limited experience with interventional EUS, they may require hands-on training with *ex vivo* or animal models followed by a standard protocol.^[14]

On the other hand, dedicated auxiliary devices can enhance the safety of EUS-GBD that are still in need before introducing the technique effectively and safely to the wider endoscopic society. In an animal study, Zhang *et al.*^[15] reported that the retrievable anchor (RA) delivered by the fine-needle aspiration needle could replace the guidewire. It is preplaced in the GB before the stent insertion, which would hold the GB close to the duodenal wall and it stabilizes the GB when introducing the stent. It is believed that the RA has several advantages. First, the anchored GB is more stable for the stent to target. Second, bile leakage may also be reduced because the puncture site is pressed by the duodenal wall. Third, in the stent release process, pulling the GB can make the internal space larger, which is more conducive to reducing the probability of the stenting failure. Of note, these anchors are nondamaging to the GB, and they are retrievable. This method heightens the probability of success by ensuring a simpler and safer procedure. Operator confidence is boosted as well.

Per-oral cholecystoscopy followed by EUS-guided gallbladder drainage

The fistula created between the GI tract and the GB enables the exploration inside the GB by the endoscope per-orally, which is also termed as peroral cholecystoscopy.^[16,17] The applications include: GB mucosa detection (*i.e.*, magnifying endoscopy observation, electronic staining, and confocal endomicroscopy examination),^[18] EUS evaluation, removing gallstones, lithotripsy of gallstones,^[19] and polyp-resection.^[20] From these, gallstone removal is the most reported therapy [Table 1].

How to perform endoscopic cholecystolithotomy Which stent used in EUS-guided gallbladder drainage can facilitate further endoscopic access?

LAMPs with an antimigration design are usually applied in the EUS-GBD, which are equipped with a "cold" or "hot" deliver system.^[23] The hot stents equipped with a cautery tipped delivery system can complete the procedure with a single-step without the tract dilation with cystotome or other dilators. This includes the HOT AXIOS stent (15 mm \times 10 mm or 10 mm × 10 mm diameter, Boston Scientific, Marlborough, Massachusetts, United States) and the Microtech stent (10 mm \times 35 mm length, Nan Jing Co. Ltd., China). The "cold" stents include the BONA-AL stent (10 mm \times 4–7 cm in length with 22 mm external flares, Standard Sci Tech Inc., Seoul, Korea) and the SPAXUS stent (10 mm × 10 mm, and $10 \text{ mm} \times 16 \text{ mm}$, Niti-S, Taewoong Medical, Korea). The overall adverse effect (AE) rate of LAMS in EUS-GBD was 13%, which includes an early AE of 6.5% and a delayed AE of 8%.[24] When deploying the LAMS, the stent-release-within-scope-channel technique was used to avoid the displacement.^[25]

What is the superior spot for the drainage: The antrum or the duodenum?

First, it should be noted that a transduodenal approach is optimal for gallstone removal in comparison to the transgastric path. If a transgastric path were chosen for

Author	Year	Cases	Spot for the drainage		Stents type	Stents	Indwelling	Stone	Spontaneous
			Stomach	Duodenum		diameter	time	clearance	stone clearance
Itoi et al.[21]	2014	1	1	N	Cold AXIOS	15 mm	14 days	1/1	Ν
Ge et al. ^[22]	2016	7	4	3	Microtech stent	10/35 mm	9 days	7/7	Ν
Larghi <i>et al</i> . ^[19]	2016	1	1	N	Hot AXIOS	15/10 mm	Ν	1/1	Ν
Chan et al. ^[18]	2017	25	6	19	Cold/hot AXIOS	15/10 mm	1-3 month(s)	22/25	14/25

Table 1.	Endoscopic	cholecyste	olithotomy
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N: Not reported.

the puncture, food impaction would occur in the GB that would be difficult to remove and it might have a potential risk of contaminating the bile duct system.^[26]

Besides, a small stone could pass through the LAMPS spontaneously when the transduodenal approach is applied. More than 10% of the cases were found with no stone residuals a couple of days after the drainage.

In a study by Chan *et al.*,^[18] food residue was found in two patients, and an attempt to prevent food entry into the GB was made by inserting pigtail stents through the LAMS, which also failed. Similar food impaction was also reported by Ge *et al.* presently, there is no effective method to prevent food impaction in the GB. As a result, drainage of the GB via the duodenum has been advocated.

When should gallstone removal be performed or does GB exploration occur after EUS-guided gallbladder drainage (the stent's indwelling time)?

In most reported studies, the indication for EUS-GBD is to treat acute cholecystitis in high-risk surgical patients.^[27] The stents should remain in place as long as possible.^[28] In a study by Choi *et al.*,^[29] the median stent patency was 190 days (15–1185 days) and 458 days (151–1185 days) for the patients who were alive at the end of the study end, and the rate of late AEs was 7%.

If the EUS-GBD was followed by a gallstone removal procedure, the path created by the stent should serve as a temperate conduit for it. The prolonged stent indwelling time after relieving the cholecystitis symptoms may not benefit per-oral cholecystoscopy. The stent should remain until the mature fistula was formed, then the endoscope could enter the GB. The rate of stent-related AEs may increase with the prolonged indwelling time. In a study by Chen,^[18] LAMS covered with granulation tissue was found in one patient, which caused the failure of a subsequent gallstone removal. The stent's indwelling time in this study was 1–3 months. In a study by Kamata *et al.*,^[6]

the metal stents were removed 4 weeks after placement in order to avoid stent migration and the recurrence of cholecystitis due to food impaction. In another study by Ligresti *et al.*,^[30] they reported that the longer stent indwelling time resulted in a buried stent and it was rescued with the LAMS-in-LAMS technique. It seems that the indwelling time is longer than 4 weeks, which may increase the risk of a stent-related AE.

The endoscopic cholecystolithotomy should be performed as soon as the symptom of cholecystitis is improved and the mature fistula is formed. In addition, the stent should be removed immediately after the successful stone removal. As reported in our previous study,^[22] the following endoscopic cholecystolithotomy was usually performed around 14 days after the EUS-GBD. On the other hand, a shorter time with <10 days of duration may not be enough for the formation of the fistulous tract and this can increase the risk of fistula rupture during the cholecystolithotomy procedure. A stent indwelling time that is <10 days may not be safe to perform peroral cholecystoscopy. However, the minimum stent indwelling time for the mature fistula formation and to facilitate the stone removal procedure with an endoscope needs to be further verified.

Procedure of stone removal

When a fistula is formed between the GI tract and the GB after cholecystostomy with ECE-LAMS, per-oral cholecystoscopy can be performed. The endoscope (e.g., gastroscope, transnasal gastroscope, or ultraslim endoscope) was advanced into the GB via the fistula formed by the stent. A stone basket was inserted into the GB to retrieve the stones [Figure 1]. The stone retrieval basket was withdrawn from the GB, and the stones were discharged into the GI tract. Water flushing may be helpful for the stone grasping procedure. After several deployments of the basket to remove the stones, an endoscope was introduced into the GB to check for any remaining stones. If the LAMS was mal-deployed and it prohibited the entry of the endoscope into the GB, or occasionally gallstones were hidden behind the

large flange of the stent, the LAMS could be removed before advancing the endoscope. If necessary, the fistula could be dilated by a balloon catheter, which can subsequently facilitate the insertion of an endoscope or extract gallstones with a larger diameter.

The device used for cholecystolithotomy

The device used for cholecystolithotomy has long been used in ERCP procedures. This is similar to holmium laser lithotripsy (SlimLine SIS GI holmium laser fiber; lumenis) for larger gallstones, fiber-optic cholangioscopy (SpyGlass, Boston Scientific), Roth net (US Endoscopy, Mentor, Ohio, USA), and a stone retrieval basket (22Q; Olympus Medical, Tokyo, Japan). Foreign body net could also be used to grasp the scattered small stones [Figure 2]. Most of these devices are designed for stone-grasping movements in a tubular lumen such as the bile duct. For example, the protruding structure at the distal end of the basket was not efficient for the GB and it may easily injure the GB mucosa while grasping for a stone within the GB. Thus, modification of the basket is necessitated so it fits into a spherical lumen.

How to guarantee stone removal without any residuals?

Cystic duct radiography is an important procedure to rule out a residual stone in the structure of the cystic duct [Figure 3]. However, injection of a contrast agent into the cystic duct with an ERCP catheter is not easy because of the existence of the Heister valve. In our experience, a lithotomy balloon can be helpful when injecting a contrast agent. There are two ways to perform cystic duct radiography. First, the balloon can be placed at the neck of the GB, and the air-filled balloon can block the proximal opening of the cystic duct, which can facilitate the contrast agent injection with pressure. Second, the balloon can be completely filled with air while within the GB and then be withdrawn until the fistula is blocked. The injected contrast agent fills the GB and goes into the cystic duct.

Follow-up after endoscopic cholecystolithotomy

The standard protocol for the follow-up after endoscopic cholecystolithotomy has not been established yet. In light of the previous experiences, we may suggest a normal diet should be resumed 48 h after the stent removal; regular ursodeoxycholic acid should be taken orally for at least 6 months; regular US examination should be performed at 1, 3, 6, and 12 months after the treatment in the 1st year and every 12 months thereafter. These procedures can be followed and verified in further studies.



Figure 1. A stone basket was inserted into the GB to retrieve the stones



Figure 2. A net was inserted to retrieve the small stones near the neck of the G



Figure 3. Cystic duct radiography to rule out a residual stone in the structure of the cystic duct

CONCLUSION

After nearly 10 years of development, EUS-GBD appears to be an attractive alternative for managing acute cholecystitis in high-risk patients. The formation of a fistulous tract after EUS-GBD facilitates the entry of the endoscope inside the GB to perform endoscopic lithotomy and polypectomy, which is an efficient and safe procedure. This technique needs to be further verified by conducting long-term multicenter prospective studies.

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Conflicts of interest

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