

Innovations in the Treatment of Gastric Outlet Obstruction: Is This the Era of Endoscopic Ultrasonography-Guided Gastroenterostomy?

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Keywords Gastric Outlet Obstruction · Endoscopic Ultrasonography-Guided Gastroenterostomy · Enteral Stent · Surgical Gastrojejunostomy

Abstract

Purpose of review Gastric outlet obstruction is a syndrome that is caused by several benign and malignant diseases. Endoscopic ultrasonography-guided gastroenterostomy (EUS-GE) is a novel treatment that seems to combine the advantages of the two traditional treatments: enteral stent placement (ES) and surgical gastrojejunostomy (SGJ). This review aims to determine the current position of EUS-GE amidst the two traditional treatment options. *Recent findings* Different approaches can be adopted to facilitate EUS-GE. Direct gastroenterostomy seems to have the shortest procedure time. All methods possess high technical success rates. Several retrospective analyses have compared EUS-GE to ES and SGJ. EUS-GE seems to have better stent patency compared to ES and a lower adverse event rate compared to SGJ. Recently, randomized trials have been initiated which compare EUS-GE with ES or SGJ. *Summary* Although the currently published literature unequivocally shows the great promises of EUS-GE, properly powered and controlled studies are warranted to ascertain the definitive position of EUS-GE within the treatment repertoire of GOO. Until reliable data has been collected, EUS-GE can be considered a safe and effective alternative to ES and SGJ.

Introduction

Gastric outlet obstruction (GOO) is a phenomenon that is characterized by mechanical obstruction of the distal stomach, pylorus, or duodenum [1]. The condition is caused by several benign and malignant diseases. Peptic ulcer disease due to nonsteroidal antiinflammatory drugs (NSAIDs) or Helicobacter pylori infection is the most frequent cause of benign GOO [2•]. Other etiologies include anastomotic strictures and (complications of) acute and chronic pancreatitis [3]. Common diseases that may result in malignant GOO are gastric, pancreatic, or duodenal carcinomas [4]. An estimated 10 to 25% of patients with pancreatic or biliary cancer will develop GOO at a certain point during the disease course [5]. Extraluminal compression of the gastric outlet by metastatic disease or lymphoma may also result in obstruction [2•].

Symptoms of GOO typically encompass nausea, vomiting, inability to tolerate oral intake, and abdominal pain [6]. Owing to metastatic or irresectable disease, patients with malignant GOO generally have a poor prognosis with a median survival of 2 to 3 months [7, 8]. These symptoms may have a considerable impact on quality of life [9]. Therefore, treatment of malignant causes of GOO is often directed at palliation of obstructive symptoms and resumption of oral intake [10, 11]. Classically, malignant GOO is treated by means of two interventions: surgical gastrojejunos-tomy (SGJ) or placement of an enteral stent (ES) (Fig. 1).

SGJ can be performed through an open or laparoscopic procedure [12]. Technical success rates of SGJ are very high, ranging from 95 to 100% [9]. Nevertheless, postoperative course is often characterized by a high morbidity rate of up to 24% [13]. In addition to delayed gastric emptying, common complications after SGJ are anastomotic leakage, ileus, and bleeding [14, 15]. These disadvantages notwithstanding, a surgically created gastrointestinal anastomosis is known to be durable in terms of patency, resulting in low reintervention rates [13].

Comparable to SGJ, ES is distinguished by a high technical success rate [16]. Moreover, postprocedural initiation of oral intake is fast, especially compared to SGJ [17]. However, ES is at risk of migration and dysfunction, due to tissue ingrowth or food impaction. Hence, reintervention because of recurrent obstructive symptoms is required in approximately 25% of patients [18].

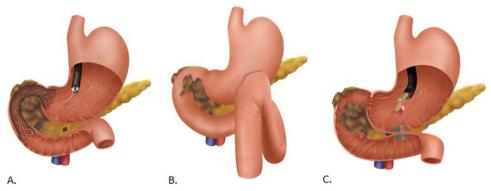


Fig. 1 Treatment options for gastric outlet obstruction. A Enteral stent placement. B Surgical gastrojejunostomy. C Endoscopic ultrasonography-guided gastroenterostomy.

Owing to the limitations of the above-mentioned treatments, endoscopic ultrasonography-guided gastroenterostomy (EUS-GE) has emerged as a potentially valuable alternative (Fig. 1) [19]. EUS-GE is characterized by placing a lumen-apposing metal stent (LAMS) between the stomach and jejunal loop distal to the

Treatment

Interventional Procedures

EUS-Guided Gastroenterostomy (EUS-GE)

obstruction, thereby creating an alternative route for food passage [20]. This minimally invasive treatment has the capacity to combine the benefits of ES and SGJ while avoiding the risks involved in both treatments [21].

This review aims to identify the current position of EUS-GE amidst SGJ and ES for the treatment of malignant GOO. Furthermore, objectives for future research will be identified. The discussion of these issues attempts to provide an answer to the question whether EUS-GE is ready for prime time.

Due to the frequently occurring negative effects of both SGJ and ES, alternative treatment options for GOO were examined in the past two decades. The first experiences with the endoscopic ultrasonography-guided creation of a gastrointestinal anastomosis by means of a specially designed suturing device were already described in 2002 [22, 23]. Nevertheless, it took about a decade for the first dedicated LAMS to be introduced [24]. The aim of LAMS placement was to create an anastomosis between two gastrointestinal lumina [25]. Initial reports focused on the creation of a cholecystogastrostomy with a biflanged stent [24]. The LAMS was loaded in a delivery system which could be connected to an echoendoscope. Originally, a guidewire was used to facilitate tract dilation and advance the LAMS into the target lumen after which the two structures could be anastomosed [24]. Updated versions of the delivery system, however, contain a wired nose cone tip that enables electrocautery. This feature promotes advancement of the catheter — in which the LAMS is folded - into the structure of interest, rendering prior tract dilation redundant [26]. Currently, LAMS placement in the gastrointestinal tract is approved for drainage of pancreatic fluid collections, the gallbladder, and biliary tract [27]. Yet, in 2012, a porcine model showed its effectiveness for the creation of a gastroenterostomy using an anchor wire, access device, and LAMS [19]. Consequently, since the first studies on EUS-GE in humans were published, this off-label indication for LAMS use gained momentum [28, 29].

Currently, two electrocautery-enhanced delivery systems are available: the Hot AXIOS[™] stent (Boston Scientific Corporation, Marlborough, MA, USA) and the Niti-S[™] Hot SPAXUS[™] stent (Taewoong Medical Co., Gyeonggi-do, South Korea) [27].

Standard Procedure

Several methods have been described to perform EUS-GE with a LAMS, such as EUS-guided balloon-occluded gastroenterostomy bypass (EPASS), balloon-assisted gastroenterostomy, and direct gastroenterostomy (synonyms: freehand technique or wireless EUS-guided gastroenterostomy simplified technique (WEST)) [30]. The balloon-assisted approach is characterized by the transgastric puncture of an inflated balloon that is positioned in the distal duodenum or proximal jejunum. Bursting of the balloon confirms that the correct target loop is punctured, after which a LAMS can be placed between the stomach and duodenum or jejunum [29]. During EPASS, a dedicated double balloon enteric tube is positioned in the distal duodenum or proximal jejunum. The balloons are inflated, and subsequently, the space between the balloons is distended by instillation of saline. Finally, this section is punctured transgastrically after which the LAMS is deployed [31, 32]. The direct gastroenterostomy approach is

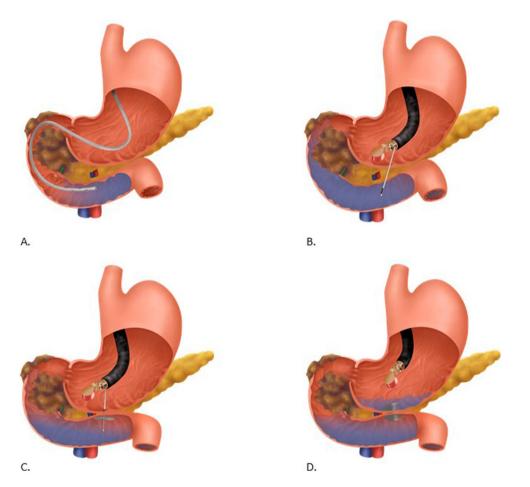


Fig. 2 EUS-GE procedural steps. A Fluid instillation of indigo-colored saline. B Transluminal puncture with electrocauteryenhanced delivery system. C Deployment of distal flange. D Deployment of proximal flange.

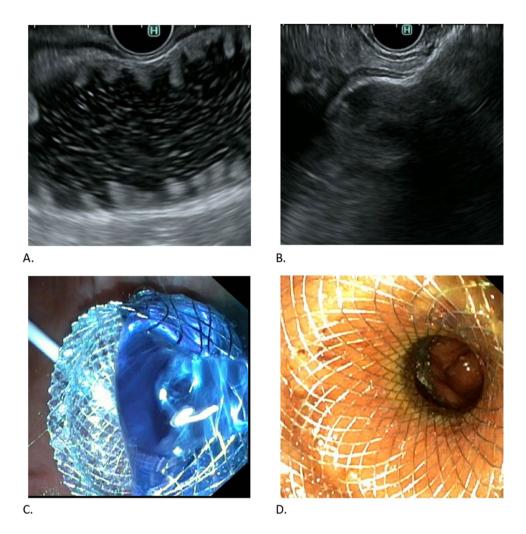


Fig. 3 Ultrasonographic and endoscopic images of EUS-GE. **A** Bowel distension after instillation of saline. **B** Deployment of distal flange in duodenum/jejunum. **C** Backflow of indigo-colored saline in stomach after deployment of proximal flange. **D** Visibility of intestinal mucosa through the LAMS.

distinguished by directly infusing saline distal to the obstruction, without using a balloon catheter or guidewire [20]. Since our institute exclusively uses this technique, we will explain the different steps involved in performing a direct gastroenterostomy in more detail (Figs. 2 and 3).

To perform EUS-GE as safe as possible, the stomach should be relieved of contents. Therefore, patients should fast for a minimum of 6 h prior to the procedure, also depending on the severity of obstruction. Preferably, a nasogastric tube is placed to minimize the risk of aspiration. Just before or after EUS-GE, prophylactic antibiotics are administered intravenously to mitigate the risk of intra-abdominal infections.

The direct gastroenterostomy is performed under deep conscious sedation or — if adequate airway management is warranted — under general anesthesia.

The first step of this approach consists in inserting a gastroscope to remove residual gastric contents. If not already in place, a jejunal feeding tube or nasobiliary drain is placed distal to the obstruction deep in the jejunum. Saline, colored with a small amount of dye (e.g., indigo carmine), is infused via the feeding tube or nasobiliary drain into the small bowel distal to the obstruction. The device we use to facilitate saline infusion is the Rapid Refill™ Continuous Injection System (Boston Scientific Corporation, Marlborough, MA, USA). At the same time as fluid is instilled, the endoscope is replaced by a therapeutic linear echoendoscope. We advise to position the echoendoscope in the gastric antrum and rotate the scope clockwise to identify a distended intestinal loop. In case of excessive intestinal motility, scopolamine butylbromide or glucagon can be used to inhibit peristalsis. Based on several endosonographic characteristics, the endoscopist is able to identify a target loop. A key feature of a saline-infused jejunal loop is what we call "the swirl sign." This entails the rapid flow of air mixed with saline close to the distal end of the jejunal feeding tube or nasobiliary drain. Another characteristic that is used is the identification of circular folds (valves of Kerckring) in the small bowel.

A target loop is sufficiently dilated when the segment is distended to at least -4 cm. This prevents that the punctured intestinal loop is pushed away from the stomach wall during LAMS deployment. Ideally, the target loop runs axially from the position of the transducer, allowing enough space to insert and deploy the LAMS. When a sufficiently distended segment of the jejunum is found, the electrocautery-enhanced delivery system can be connected to the echoendoscope. If the endoscopist is unsure whether the target loop is in fact jejunum, it is recommended to puncture the lumen with a fine needle aspiration (FNA) needle and verify that the punctate is indeed blue dyed saline. This confirms that a correct loop is visualized.

After verification of the correct target lumen, the stomach wall and intestinal wall are punctured consecutively with the electrocautery-enhanced catheter. Preferentially, the stomach is punctured as distal as possible. The distal flange of the 20-mm LAMS is deployed in the jejunum followed by traction of the flange against the bowel wall. Subsequently, the proximal part of the stent is deployed in the stomach and pushed out the working channel during gentle retraction of the echoendoscope. Position and passage of the stent are confirmed by backflow of the indigo-colored saline from the jejunum back into the stomach. The saline will be suctioned to prevent aspiration pneumonia. We generally do not dilate the LAMS; instead, it will be left to expand naturally. In case EUS-GE is used to palliate symptoms of malignant GOO, the LAMS will not be removed. Directly after a technically successful procedure, all tubes will be removed.

Patients are allowed a clear liquid diet the same day as the procedure took place. The next morning, this can be expanded to an easily digestible diet if there are no signs of complications. When the patient tolerates this without vomiting, oral intake can be resumed without any restrictions. Generally, patients are discharged the day after the procedure, although limited data suggests that same-day discharge may be safe as well. No prospective comparisons of the different EUS-GE techniques have been conducted. Chen et al. retrospectively compared the direct and balloon-assisted techniques. They concluded that the direct method could be performed in less time than the balloon-assisted approach. However, no differences in technical success, clinical success, and adverse events (AEs) were found [33]. A recently published meta-analysis, on the other hand, did find a lower AE rate in the direct EUS-GE group as compared to balloon-assisted techniques. The same applies to the length of hospital stay [30]. Primarily based on the suggestion that the direct approach consumes less

time, many endoscopists prefer this method [20].

General Outcomes of EUS-GE

To date, more than 20 single-arm studies have been published that specifically focused on EUS-GE performance and outcomes in malignant GOO. Most have a sample size between 10 and approximately 50 patients, with few exceeding 100 patients. Based on these studies, technical and clinical success ranges between 80–100% and 73–95%, respectively [21, 34, 35, 36•].

A meta-analysis from Iqbal et al. reported an AE rate of 12% after EUS-GE [37]. Serious AEs occur in approximately 3–6% of EUS-GE procedures, consisting of peritonitis, perforation, hemorrhage, and abdominal pain [38, 39]. A notorious periprocedural complication of EUS-GE is LAMS misdeployment. This occurs when the distal or proximal flange of the LAMS is deployed outside of the gastrointestinal tract. When the target lumen is already punctured, this might have serious consequences, such as peritonitis, surgical intervention, or death [40•]. Although misdeployment is not uncommon — occurring in up to 10% of LAMS placements — most of these complications can be managed endoscopically and have only mild consequences [40•]. The rate of LAMS misdeployments might be related to the technical complexity of EUS-GE and the learning curve associated with the procedure [40•, 41•, 42•]. It is estimated that proficiency is achieved after 7 to 25 procedures. Notably, the studies that have assessed learning curve of EUS-GE included endoscopists experienced in therapeutic EUS, emphasizing the complexities associated with the procedure [41•, 42•].

Currently, LAMSs are registered for implantation up to a maximum of 60 days in case of drainage of pancreatic fluid collections [43]. However, in case of palliative treatment of GOO, the LAMS generally remains in situ until death or until removal is warranted for medical reasons. A long-term evaluation of patients who underwent EUS-GE observed LAMS dysfunction in 2 of 53 patients (3.7%) because of stent occlusion. The other patients experienced no LAMS-related problems, after a median follow-up of 196 days in malignant GOO and 319.5 days in benign GOO [44]. These results are corroborated by Kastelijn et al., who observed a LAMS dysfunction rate of 6.1% [21].

In addition to LAMS dysfunction as a long-term complication of EUS-GE, some case reports have described jejunal ulceration at the site opposite to the distal flange, even resulting in perforation and consequent peritonitis. Yet, this

| | z | | EUS-GE method | Technical success (%) | : (%) Clinical success (%) | cess (%) | Adverse events (%) | ents (%) | LAMS |
|--------------------------|-----------------|-----------------|-----------------------|-----------------------|----------------------------|----------|--------------------|----------|--------------------------|
| | | | | | | | | | misdeployment (N (%)) |
| | EUS-GE | ß | | EUS-GE | ES EUS-GE | ន | EUS-GE | ß | I |
| Chen (2017) | 30 | 52 | Various** | 86.7 | 94.2 83.3 | 67.3 | 16.7 | 11.5 | 3 (10.0) |
| Ge (2019) | 22 | 78 | Direct | 100.0 | $100.0 95.8^*$ | 76.3 | 20.8 | 40.2 | 2 (9.0) |
| Van Wanrooij (2022) | 88 [†] | 88 [†] | Direct | 94.0 | $98.0 \ 91.0^*$ | 75.0 | 10.0 | 21.0 | 3 (3.4) |
| Sánchez-Aldehuelo (2022) | 79 | 97 | Direct | 93.7 | 92.8 92.4 | 83.5 | 10.1 | 10.3 | 3 (3.8) |
| Chan (2022) | 30 | 49 | EPASS | 93.3 | 100.0 93.3 | 87.8 | 6.7 | 26.5 | 1 (3.3) |
| Jaruvongvanich (2023) | 232 | 131 | Various ^{**} | 98.3 | $99.2 98.3^*$ | 91.6 | 8.6^* | 38.9 | 4 (1.7) |

| | z | | EUS-GE method Technical success (%) Clinical success (%) Adverse events (%) LAMS misde (*, (%) | Technical | success (%) | Clinical s | uccess (%) | Adverse e | /ents (%) | LAMS misdeployment (1, 1,02,1) |
|---------------------------|-----------------|-----|--|-----------|-------------|------------|------------|------------|-----------|--------------------------------------|
| | EUS-GE SGJ | SGJ | | EUS-GE | SGJ | EUS-GE | 2GJ | EUS-GE | SGJ | |
| Khashab (2017) | 30 | 63 | Balloon-assisted | 87.0 | 100.0^{*} | 87.0 | 90.06 | 16.0 | 25.0 | 3 (10.0) |
| Perez-Miranda (2017) | 25 | 29 | Various** | 88.0 | 100.0 | 84.0 | 90.0 | 12.0^{*} | 41.0 | 9 (36.0) |
| Bronswijk (2021) | 37 [†] | 37† | Direct | 94.6 | 100.0 | 97.1 | 89.2 | 2.7* | 27.0 | 2 (5.4) |
| Kouanda (2021) | 40 | 26 | Direct | 92.5 | 100.0 | 85.0 | 84.0 | 22.5 | 88.5 | 1 (2.5) |
| Abbas (2022) | 25 | 27 | Direct | 100.0 | 100.0 | 88.0 | 85.0 | 8.0* | 41.0 | 1 (4.0) |
| Chan (2022) | 30 | 35 | EPASS | 93.3 | 100.0 | 93.3 | 80.0 | 6.7 | 28.6 | 1 (3.3) |
| Jaruvongvanich (2023) 232 | 232 | 73 | Various** | 98.3 | 100.0 | 98.3^{*} | 91.6 | 8.6^{*} | 27.4 | 4 (1.7) |

The Era of Endoscopic Ultrasonography-Guided Gastroenterostomy

| | EUS-GE | B | SGJ |
|---------------|---|---|--|
| Advantages | Minimally invasive Fast resumption of oral intake High technical success rate | Minimally invasive Fast resumption of oral intake High technical success rate | Very high technical success rate Durable anastomosis |
| | (Potentially) durable anastomosis Relatively small risk of adverse events Short hospital stay | Safe procedure Short hospital stay | High long-term patency |
| Disadvantages | • • | High risk of stent obstruction or migration and sub- sequent reinterventions and need for hospitalization | Considerable postoperative morbidity Larger interval between proce- |
| | potentially fatal consequences Uncertainty on the long-term patency | | dure and clinical successLonger hospital stay |

Table 3. Advantages and disadvantages of EUS-GE, ES, and SGJ

complication is perhaps more common in benign GOO, since indwelling time tends to be longer compared to malignant causes of GOO [45].

Comparison with ES

With regard to the effectiveness of EUS-GE, several retrospective cohort studies have compared LAMS placement with ES (Tables 1 and 3) [46••, 47••, 48••, 49••, 50••, 51••]. None of these studies observed any differences in technical success. However, three studies concluded that EUS-GE resulted in higher clinical success rates compared to ES. These studies observed clinical success in more than 90% of EUS-GE patients, whereas ES was clinically successful in approximately 75% of patients [46••, 48••, 51••].

Only the study of Jaruvongvanich et al. reported statistically significant differences in the occurrence of AEs, in favor of EUS-GE (8.6% vs. 38.9%) [51••]. With regard to the patency of both treatments, four studies observed a significantly lower reintervention rate for recurrent obstructive symptoms in patients who underwent EUS-GE [46••, 47••, 48••, 51••]. This discrepancy is primarily explained by the high rate of stent dysfunction due to tumor in- or overgrowth, food impaction, or migration after ES. Moreover, time to stent failure also seems to be shorter after ES, compared to EUS-GE [48••, 49••, 51••]. With regard to EUS-GE, the pooled rate of LAMS misdeployment of these cohorts combined was 3.3%. Partly based on these comparative studies, the Endoscopic Society of Gastrointestinal Endoscopy (ESGE) currently perceives the use of EUS-GE in an expert setting as a valid alternative to ES [52•]. Nevertheless, the results of these retrospective studies should be interpreted cautiously since they might be affected by selection bias.

Comparison with SGJ

EUS-GE has been compared to SGJ in seven retrospective studies (Tables 2 and 3) [50••, 51••, 53••, 54••, 55••, 56••, 57••]. In all studies, technical success of SGJ was 100%. Most studies did not find differences in technical success between the two approaches. However, Khashab et al. observed a higher technical success rate when SGJ was performed (100% vs. 87%), although this did not translate into a higher proportion of clinical success [57••]. This is comparable to most other studies [50••, 53••, 54••, 55••, 56••, 57••]. The largest study of Jaruvongvanich et al. did find differences in clinical success — defined as tolerance of at least a liquid diet within 2 weeks — favoring EUS-GE (98.3% vs. 91.6%). Moreover, this study found a significantly lower reintervention rate in the EUS-GE group (0.9% vs. 13.7%) [51••].

Four studies noted a lower AE rate after EUS-GE, compared to SGJ. AEs in patients who underwent SGJ mainly consisted of postoperative delayed gastric emptying or ileus, anastomotic leakage, bleeding, or infection [51••, 53••, 54••, 56••]. Remarkably, Abbas et al. observed a lower AE rate after EUS-GE compared to SGJ in patients with peritoneal carcinomatosis, even though the former group was in a clinically worse condition than patients who underwent SGJ [56••].

What should be noted is that some studies compared EUS-GE with an open surgical approach, in contrast to a laparoscopic approach, which could have influenced the results [55••, 56••, 57••]. A laparoscopic gastrojejunostomy is considered to be the better option, since it results in a lower rate of postoperative gastroparesis, faster clinical success, and shorter length of hospital stay [58].

Bronswijk et al. and Kouanda et al. observed a shorter time to oral intake after EUS-GE (median 1 vs. 3 days and mean 1.3 vs. 4.7 days, respectively) [54••, 55••]. Moreover, time to initiation or resumption of chemotherapy tends to be shorter after EUS-GE compared to SGJ [55••, 56••].

Largely based on the above-mentioned publications, the ESGE concludes that EUS-GE is a valid alternative to SGJ within an expert setting [52•]. However, similarly to the comparison of EUS-GE with ES, the results of these studies might be affected by selection bias.

Practical Considerations

Based on our personal experience with EUS-GE and supported by evidence from the literature, we have formulated the following practical considerations which can be taken into account when considering EUS-GE:

We recommend the use of a LAMS with the *largest luminal diameter*, as to ensure an as wide as possible gastrointestinal anastomosis. Currently, the AXIOS[™] stent has a maximum available diameter of 20 mm, whereas the SPAXUS[™] stent has a largest available diameter of 16 mm [59, 60]. A recently published retrospective analysis compared the use of the 15 mm AXIOS[™] stent with the 20 mm stent. Although the authors did not observe differences in clinical success and the rate of AEs, significantly more patients with a 20-mm LAMS tolerated a soft solid or complete diet at the end of study follow-up [36•]. These results suggest that a LAMS with the largest diameter is preferred.

The current literature did not specifically focus on the relation between *etiology of the obstruction* and technical and clinical success. Based on experience in our institute, we speculate that gastric cancer as cause of GOO might negatively influence EUS-GE performance and efficacy. The presence of a malignancy at the region of the antrum or pylorus could result in an increased distance between the gastric wall and the jejunal target loop, thereby creating difficulties for safe puncture and deployment of the LAMS. Moreover, malignancies located in the distal stomach might decrease gastric motility, which could lead to disappointing clinical effectiveness of EUS-GE. Currently, our research group is assessing the influence of etiology on EUS-GE outcomes. Hence, a more definitive conclusion is expected in time.

Malignant GOO may occur in conjunction with *biliary obstruction* [61]. Limited evidence is available with regard to the preferred treatment options in these specific situations. The CABRIOLET-study evaluated the relationship between various treatment combinations (EUS-guided interventions,

transpapillary and/or duodenal SEMS placement) and dysfunction rates [62•]. Although highly underpowered, the results suggest a preference of EUS-GE with EUS-guided hepaticogastrostomy over other combinations, such as EUS-GE with transpapillary stent placement, EUS-guided cholecys-toduodenostomy (EUS-CDS), or ES combined with either one of the above-mentioned procedures. The former treatment combination may lead to the lowest recurrence rate. The use of EUS-CDS, on the other hand, seems to be related to worse outcomes since it may be prone to food impaction due to a "sump" that is created when the obstructed duodenal bulb becomes clogged with food remains [62•]. Despite the drawbacks of this retrospective analysis, these outcomes suggest that certain treatment combinations might be better suited for concomitant biliary and gastric outlet obstruction than others.

Peritoneal involvement is predictive of morbidity and mortality in patients undergoing SGJ for malignant GOO [63•]. One retrospective study assessed whether this was also the case for EUS-GE compared to SGJ [56••]. Albeit EUS-GE patients were older and had more advanced disease, there was no difference in technical success. Furthermore, EUS-GE was associated with a lower AE rate. What should be noted is that patients in the SGJ group underwent an open procedure, what may have influenced the results. Nevertheless, in our experience, EUS-GE in patients with peritoneal carcinomatosis is generally safe and effective. However, it is advisable to confirm adequate intestinal passage by placing a nasojejunal feeding tube in order to exclude distal obstruction and paralytic ileus. Moreover, if abdominal radiology shows diffuse dilation of the small bowel with air-fluid levels, we do not recommend EUS-GE since risk of clinical failure tends to be high.

The presence of *ascites* is considered a contraindication for save execution of EUS-GE [52•]. This has various reasons. First, diagnosing ascites often implies the presence of peritoneal metastases and hence an advanced disease stage, rendering the clinical benefit of EUS-GE less likely [52•, 64•]. Second, ascites might suggest the existence of distal obstruction sites due to peritoneal metastases [52•]. Third, ascites might pose technical challenges during LAMS placement since it could push the target loop away from the gastric wall [64•]. Lastly, the electrocautery-enhanced tip of the delivery system may function less optimally. Based on these potential problems, we advise prudence when patients with GOO and ascites are considered for EUS-GE. Nevertheless, a retrospective analysis from Basha et al. compared patients with and without ascites that underwent EUS-GE, and they did not find differences in technical and clinical success or in the rate of adverse events [64•]. Hence, if the space between the gastric wall and the target loop is not increased by the presence of ascites, we argue that one can proceed with EUS-GE.

Despite LAMS misdeployment being a relatively uncommon complication of EUS-GE, consequences might be dreadful [40•]. In many cases, however, serious consequences can be prevented by merely placing through-the-scope or over-the-scope clips or perform endoscopic suturing [40•]. Nevertheless, a systematic review identified several methods to endoscopically salvage a misdeployed LAMS, such as bridging the initially placed LAMS with a second LAMS or SEMS [65]. An alternative approach is natural orifice transluminal endoscopic surgery (NOTES), in which the peritoneal cavity is traversed through the original gastric puncture site, a jejunal loop is grasped and penetrated with a needle-knife, allowing a second stent to be placed in the target jejunal lumen [66]. The above-mentioned salvage methods require considerable endoscopic expertise. Therefore, it might be desirable to be in close contact with the surgeon on call when performing EUS-GE, facilitating rapid discussion regarding the preferred way to treat the complication.

Accidental creation of a *gastrocolic anastomosis* is rare, occurring in less than 5% of EUS-GE misdeployments [40•]. In our institute, we have experienced this complication in one patient. She was admitted with post-prandial diarrhea, which led to the radiologic confirmation of the misplaced LAMS about a month after initial placement. We believe the risk of creating a gastrocolostomy increases when the interval between saline infusion and actual LAMS placement is prolonged, since the instilled fluid will not only dilate the jejunum but eventually the colon as well. Endoscopic ultrasound may not always be feasible to adequately differentiate a large bowel loop from a small bowel loop. Hence, caution is advised when identification of a target loop appears to be complicated and risk of saline infusion in the colon increases.

EUS-GE is primarily described in light of malignant GOO. However, a limited number of retrospective studies specifically assessed EUS-GE as treatment for *benign* GOO, demonstrating excellent results in terms of technical success [3, 67]. However, in contrast to patients in a palliative setting, the LAMS is expected to be indwelling for a longer duration. This poses additional concerns, such as bleeding or a buried LAMS [3]. James et al., however, observed a relatively low rate of surgical reinterventions (16.6%) in the period prior to resolution of the obstruction, suggesting that the LAMS can safely remain in situ until the cause of GOO is definitively solved. These results are based on small sample sizes, and therefore, we recommend leaving the LAMS indwelling for a minimum amount of time, thus decreasing the risks of long-term complications.

Future Directions

A steadily growing collection of evidence already has EUS-GE firmly embedded in the treatment arsenal of GOO. Numerous hospitals worldwide have already adopted EUS-GE as the primary treatment of choice of this debilitating syndrome [62•]. Nevertheless, up until today, LAMSs are not CE-marked or FDA-approved for creating a gastrointestinal anastomosis [27]. Therefore, future research should concentrate on outcomes that are sincerely meaningful from a patient perspective. In our opinion, two aspects are of special interest. First, short-term outcomes of EUS-GE, such as time to resumption of oral intake and length of hospital stay, should be assessed prospectively. Second, studies ought to focus on long-term LAMS patency, e.g., the rate of recurrent symptoms or reinterventions. Preferably, these studies should have a comparative design. Currently, several randomized controlled trials (RCTs) are planned or have recently started patient accrual. These results are eagerly awaited. Two studies compare EUS-GE with ES and are powered on the rate of GOO recurrence and reinterventions [68, 69]. Four other studies randomize patients between EUS-GE and SGJ and focus on either functional recovery, time to oral intake, or adverse events [70–73]. Moreover, our institute initiated the Dutch ENDURO-study. This RCT is unique in the sense that it is powered on two coprimary endpoints, focusing on short-term efficacy (time to oral intake of at least soft solids) and longer-term durability (persistent or recurrent symptoms of GOO requiring reintervention) [74]. In the end, the evidence gathered by these trials might be sufficient to definitively determine the place of EUS-GE for the treatment of GOO.

Conclusions

EUS-GE is an exciting and promising new approach for the treatment of GOO. It may combine the advantages of ES and SGJ, without adopting the negative effects. However, it should be emphasized that the currently published retrospective literature is prone to selection bias, rendering appropriate comparison between EUS-GE and the other existing treatments of GOO difficult. ES might still be the treatment of choice for patients with an expected short survival due to the small interval between treatment and clinical success. When comparing SGJ to EUS-GE, the latter might have a more beneficial risk profile. Nevertheless, EUS-GE remains a high-risk procedure which requires considerable expertise. Moreover, long-term patency of the LAMS has not been sufficiently compared with the patency of a surgical bypass. Therefore, well-designed trials are necessary to definitively determine the place of EUS-GE amongst ES and SGJ. Steadily, more and more hospitals with expertise in advanced endoscopy have adopted the treatment. Until reliable data has been collected, EUS-GE can be considered a safe and effective alternative to ES and SGL

Acknowledgements

We are grateful to ClaudiaStreken for creating the illustrations in this review (Fig. 1 and Fig. 2).

Declarations

Conflict of Interest

Yorick van de Pavert reports no conflicts of interest. Leon Moons is a consultant for Boston Scientific. Auke Bogte reports no conflicts of interest. Frank Vleggaar is a consultant for Boston Scientific.

Human and Animal Rights and Informed Consent

This article does not contain any studies with human or animal subjects performed by any of the authors.

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