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Upper Gastrointestinal Stent Insertion in Malignant and Benign Disorders

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Upper gastrointestinal (GI) stents are increasingly being used to manage upper GI obstructions. Initially developed for palliative treatment of esophageal cancer, upper GI stents now play an emerging role in benign strictures of the upper GI tract. Because recurrent obstruction and stent-related complications are common, new modifications of stents have been implemented. Self-expandable metal stents (SEMS) have replaced older plastic stents. In addition, newly designed SEMS have been developed to prevent complications. This review provides an overview of the various types, indications, methods, complications, and clinical outcomes of upper GI stents in a number of malignant and benign disorders dividing the esophagus and gastroduodenum.

Key Words: Upper gastrointestinal tract; Stents; Obstruction; Malignant; Benign

INTRODUCTION

Upper gastrointestinal (GI) stents are devices used to maintain bowel luminal patency, thereby increasing nutrition intake and improving quality of life.^{1,2} Upper GI stents have mainly been developed to alleviate malignant obstructions caused by esophageal cancer, gastric cancer, periampullary cancer, or other malignancies that directly invade the upper GI tract. Disadvantages of GI stents are recurrent obstruction and stentrelated complications. In the last decade, studies in this area have focused mostly on modifying stent design to improve efficacy and safety. In addition, indications for upper GI stents have widened to include benign diseases, such as benign stricture and perforation. However, most studies in this area are small and uncontrolled. The optimal indications, methods, and duration of therapy for these disorders remain to be determined.

This review provides an overview of upper GI stenting in the esophagus and gastroduodenum, with an emphasis on the

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indications, methods, complications, and clinical outcomes.

TYPES OF STENTS

Upper GI stents were originally designed as rigid cylinders, resulting in poor efficacy and high complication rates. In recent decades, several types of flexible and self-expandable stents have been developed. Currently available upper GI stents include self-expandable metal stents (SEMS) for esophageal and gastroduodenal obstructions and self-expandable plastic stents (SEPS) for esophageal obstructions.³ Metal stents, made of stainless steel and alloys such as nitinol and Elgiloy (Elgiloy Specialty Metals, Elgin, IL, USA), have a higher degree of flexibility and are capable of generating high radial forces to maintain stent patency and position. Nitinol is an alloy of nickel and titanium and is most commonly used as a material for SEMS. It possesses good shape-memory properties and flexibility.⁴ SEMS are available as uncovered, partially covered, or fully covered with a coating, usually a plastic membrane or silicone.3 SEPS are composed of polyester and silicone and are indicated for the treatment of malignant esophageal obstructions. Although there were no differences in the clinical outcomes using SEPS and SEMS in terms of symptom relief, complications, or survival, the insertion of SEPS is technically more difficult, with a higher migration rate than that of SEMS in esophageal malignant obstructions.^{5,6} Each type

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of stent has its own inherent optimal indications, advantages, and disadvantages in clinical situations (Table 1).

Stent innovations include antireflux and antimigration features. Antireflux features were especially developed for stents through the lower esophageal sphincter. Generally, a one-way valve was attached to the distal end of the stent, inhibiting reflux from gastric contents into the esophagus. Although some studies indicated that reflux symptoms improved with the use of anti-reflux stents, a meta-analysis did not identify any significant differences in reflux symptoms, complications, or quality of life.7 Moreover, as this type of stent is covered, it migrates more frequently. Therefore, the use of anti-reflux stents has been all but abandoned. Antimigration stents have properties such as a metal mesh, wider diameter with stent flares, full double-layered coverage, and antimigration rings on the outer surface, such as anchoring devices. However, several studies did not find any significant differences in migration.89 Moreover, frequent recurrent obstruction and complications were reported.10,11

ESOPHAGEAL STENTS

Indications

The goals of stent placement are to provide relief from obstructive symptoms, allow the patient to resume a diet, and improve the patient's quality of life. As such, stent placement is not indicated for asymptomatic patients with upper GI obstructions who can tolerate a normal diet. Other contraindications to stent placement are the presence of multiple bowel obstruction sites (since stenting the proximal obstruction is unlikely to provide symptom improvement) and the presence of free perforations.

The indications for esophageal stents include malignant or

benign diseases in the esophagus, gastroesophageal junction, and gastric cardia. The earliest indication for a GI stent was a malignant obstruction by esophageal cancer and gastric cardia cancer. Extrinsic tumor invasion or compression may also be another indication. Recently, the use of esophageal stents has increased to include benign strictures (peptic, caustic, and radiation-induced), postoperative leaks, iatrogenic perforations, tracheoesophageal fistulas, and refractory esophageal variceal bleeding. Palliative care of inoperable malignant obstruction remains the most common indication for esophageal stenting (Table 2).

Methods

During stent selection for esophageal obstruction, the location of the obstruction, length of stent, and type of stent should be considered. For upper obstructions, the use of esophageal stents is considered to relatively contraindicative due to the risk of intolerable foreign body sensations, pulmonary aspiration, and migration into the hypopharynx. Modified stents with a shorter flange and an obtuse angle may be beneficial in preventing cervical esophageal obstruction.^{12,13} Due shortening of stents after deployment, it is important to select a stent length of at least 4 cm longer than the length of the stricture. This increases the likelihood of an adequate margin at the proximal and distal ends of the obstruction.

Given the potential risk of airway compression with stent expansion, esophageal stent insertion is frequently performed via endoscopy under sedation. Standard procedural sedation with an opiate and benzodiazepine is usually sufficient. The monitoring of blood pressure and oxygen saturation is important, and the patient should be placed in the left lateral decubitus or prone position to minimize the risk of aspiration due to retained gastric contents. The prone position allows for

Table 1	١.	Comparison	of the	Indications,	Advantages,	and Disadvantage	es of Different	Types of Stents
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Types of stent	Appropriate indications	Advantages	Disadvantage
Uncovered SEMS	Expected poor survival	Low risk of migration	High risk of tumor ingrowth
	High risk of migration		Difficult removal
	Gastroduodenal obstruction to avoid		
	obstructing biliary drainage		
Partially covered SEMS	High risk of migration in patient otherwise suitable for fully covered SEMS	Intermediate risk of migration and tumor ingrowth	Intermediate risk of tumor ingrowth Difficult removal
Fully covered	High risk of tumor ingrowth	Low risk of tumor ingrowth	High risk of migration
SEMS	Temporary measure: benign stricture	Easy removal	
SEPS	Benign esophageal stricture	Safe and easy removal	High risk of migration
	Temporary measure	No tumor ingrowth	Complex and stiff stent introducer system
			High rate of stent failure requiring reintervention

SEMS, self-expandable metal stent; SEPS, self-expandable plastic stent.

Site	Benign disorders	Malignant disorders
Esophagus	Benign stricture	Esophageal cancer
	Esophageal perforation	Gastric cardia cancer
	Esophagobronchial fistula	Extrinsic tumor invasion or compression such as lung cancer, mediastinal
	Refractory esophageal	cancer, metastatic cancer
	Variceal bleeding	
Gastroduodenum	Benign stricture (possible)	Gastric cancer
		Duodenal cancer
		Pancreatic cancer

Table 2. Indication for Stent Insertion in the Upper Gastrointestinal Tract

a better anatomic view under fluoroscopy. The supine position should be avoided unless the patient is endotracheally intubated due to the risk of aspiration. First, the length of stenosis should be estimated by endoscopy, fluoroscopy, or computed tomography prior to stent insertion. When estimation of the length of stenosis is impossible, a stent long enough to provide complete coverage of the full stricture should be selected. Second, a guide-wire is inserted through the obstructive lesion, with or without fluoroscopic guidance, at least 20 cm distal to the obstruction.^{12,14} Adequate insertion of the guide-wire can be confirmed by fluoroscopy and loss of resistance to the guide-wire. Third, the stent is inserted through the guide-wire up to roughly 2 cm proximal to the obstructive lesion. To assist adequate stent deployment, the proximal and distal ends of the stent are marked accordingly with radiopaque markers. Finally, the delivery device is withdrawn with endoscopic guidance, maintaining the stent in place. Complete stent expansion usually occurs within 24 to 48 hours, although very tight strictures may lead to longer or incomplete expansion. Patients are permitted to ingest liquids following the procedure and, subsequently, cautiously advance to a low-residue diet.¹⁵ The diet can start gradually from water to a regular diet within 24 to 48 hours following stent insertion.⁴ Afterward, simple radiography should be used to confirm proper location and expansion of the stent.

Complications

Potential stent-related complications include recurrent obstruction due to tumor ingrowth, perforation, stent migration, bronchoesophageal fistulas, and retrosternal pain.

Recurrent obstruction occurs in almost 30% of patients and develops most often due to tumor ingrowth and food impaction. In case of tumor ingrowth, insertion of a second stent is effective in restoring luminal patency.^{16,17} This complication can also result from stent migration. Endoscopic repositioning or reinsertion of a new stent may be preferable. Obstruction by impacted food can be managed by endoscopic clearance. Another rare late complication is stent fracture, resulting in recurrent obstruction. In this situation, stent-in-stent insertion appears to be safe and effective. Endoscopic removal of fractured stents is also possible.¹⁷

Migration is more common for covered stents than uncovered stents. The reported rate of migration is 10% to 25% for covered stents and 2% to 5% for uncovered stents. Migration can reportedly be prevented by clipping of the proximal end to the esophageal wall.¹⁸

Iatrogenic esophageal perforations are life-threatening, with a high rate of morbidity and mortality. Surgical treatment for these patients is associated with poor outcomes.¹⁹ Stent-associated esophagobronchial fistulas occurs most frequently several months after stent insertion. A recent retrospective study reported that stent-associated esophagobronchial fistulas developed in 4% of patients with esophageal stenting.²⁰ Additional insertion of a fully covered SEMS is an effective treatment modality in this case.

Retrosternal pain occurs in 13% to 60% of patients.^{2.21} Pain continues for an average of 10 days, becoming tolerable in most patients with the administration of analgesics without the need for stent removal.

Clinical outcome

Over the last 15 years, six randomized trials comparing different stents in patients with malignant esophageal obstructions have been published.²²⁻²⁷ Technical success is defined as successful insertion and adequate placement of the stent. Clinical success is the palliation of dysphagia. The technical and clinical success rates in these studies were 83% to 100% and 80% to 95%, respectively. In previous studies, the rate of recurrent dysphagia was reported to be 8% to 52%. The median duration of esophageal stent patency was reported to be 94% at 4 weeks, 78% at 3 months, and 67% at 6 months. Major complications, including perforation, aspiration pneumonia, stridor, and overt hemorrhaging, were reported for 8% to 36% of cases. A recent study showed that insertion of a novel esophageal stent coated with 125I seeds provided relief from dysphagia and prolonged survival compared to a conventional covered SEMS in patients with unresectable esophageal cancer.27

In benign strictures, the mainstay of management has been endoscopic balloon dilation.²⁸ Efficacy varies depending on the etiology of the strictures and clinical improvement in anastomotic, caustic, peptic, and radiation-induced strictures has been reported to be 92%, 84%, 81%, and 58%, respectively.²⁹ However, patients with complex strictures have a high risk of persistent or recurrent strictures. The concept of an esophageal stent in refractory strictures depends on continuous dilation of the stricture site for at least 6 weeks, causing sustained luminal patency after stent removal. Various types of stents have been explored for this indication, including extractable SEMS, SEPS, and biodegradable stents. Overall, sustained improvement of dysphagia was achieved in approximately 50% of patients.30 Long-term efficacy of SEPS was not found to be different compared with SEMS in terms of relief of dysphagia, although migration rate was higher. In the largest prospective study, stent migrations were observed in 22% of patients.³¹ Moreover, SEPS were associated with frequent complications, including severe chest pain, perforation, fistulas, and bleeding.^{32,33} Biodegradable stents do not require removal. Several studies have evaluated these stents in refractory benign strictures.³⁴⁻³⁶ In the largest study, recurrent dysphagia occurred in 21 of 28 patients at a median of 90 days and was caused by recurrent strictures, stent migration, and food impaction.³⁷ Esophageal perforations are life-threatening, with a high rate of morbidity and mortality.³⁸ Prompt intervention is therefore very important in the treatment of these patients. Insertion of a covered stent sealing the perforation site has evolved as a promising treatment modality. A systemic review showed perforation healing in 85% of cases,19 with an overall mortality of 13%, lower than reported in surgical treatment. Refractory esophageal variceal bleeding after a failed endoscopic variceal ligation is another interesting indication for esophageal SEMS insertion. The efficacy and safety of specially designed stents (SX-Ella stent Danis; Ella-CS, Hradec Kralove, Czech Republic) has been evaluated in several case series.³⁹⁻⁴² Hemostasis was achieved in 77% to 100% of patients. Stent migration has been reported in up to 25% of cases, though endoscopic repositioning was feasible. These results suggest that stent insertion can be used in refractory esophageal variceal bleeding.

GASTRODUODENAL STENTS

Indications

Gastroduodenal obstruction causing gastric outlet obstruction is common in patients with malignancies of the stomach, duodenum, and pancreas. In the Western world, malignant gastric outlet obstruction occurs most commonly due to obstruction by pancreatic cancer. Malignant gastric obstructions at the level of the stomach due to stomach cancer is seen in Asian countries, including Korea.43 These patients have nausea, vomiting, early satiety, weight loss, and a poor quality of life. Gastroduodenal stent placement is used to treat symptomatic patients with unresectable or recurrent malignancies at anastomotic sites. Patients considered for stent placement should have a short life expectancy (less than 2 to 6 months).^{44,45} As with other enteral stents, stent insertion is not indicated for asymptomatic patients with a gastric outlet obstruction and who tolerate a regular diet. Other contraindications to stent placement are the presence of multiple small bowel obstruction sites, since stenting the proximal stricture is unlikely to provide symptom relief, and the presence of free perforations. In addition, patients with peritoneal carcinomatosis may not respond as well to stent placement due to the risk of multiple obstruction sites, although some data suggest that stents can work in such patients. In a retrospective study of 215 patients undergoing stent placement for malignant gastric outlet obstruction, 116 patients (60%) had carcinomatosis and 76 (40%) did not.⁴⁶ Clinically successful outcomes (81% vs. 84%, respectively), need for reintervention (18% vs. 27%, respectively), and major complications (4% for both groups) were similar for those with and without carcinomatosis.

There have been several reports on gastroduodenal stenting in patients with benign gastroduodenal obstructions who have high risks in surgery.⁴⁷⁻⁴⁹ However, it seems premature to consider stent insertion as an alternative therapeutic modality for surgery or endoscopic balloon dilation due to frequent migration (Table 2).

Methods

It may be important to perform a radiographic study with computed tomography with oral contrast or an upper GI series to assess the anatomy, length of stricture, and degree of obstruction, especially in patients where it is unclear if the obstruction is functional.⁵⁰ Other techniques are the same as with esophageal stents.

Complications

Several complications can occur during or after gastroduodenal stent insertion.^{51,52} Intraprocedural complications include those related to sedation, pulmonary aspiration, stent malposition, perforation, and bleeding. Late complications include stent migration, stent occlusion, fistula formation, perforation, bleeding, and occlusion of biliary stents.

In a systematic review of 606 patients, there was no procedure-related mortality. Severe adverse events, including bleeding and perforation, were observed in seven patients (1.2%), stent migration in 31 patients (5%), and stent obstruction in 104 cases (18%), mainly due to tumor ingrowth. Obstruction of stents by tumor ingrowth of malignant or benign tissue is usually managed by the placement of additional stents through the original ones.⁵³ There are currently no data concerning the safety of gastroduodenal stenting in patients who have undergone or are undergoing radiation

Patients with a SEMS for a biliary obstruction who subsequently undergo duodenal stent placement are at an increased risk of biliary stent dysfunction. In a series of 410 patients with biliary stents, 33 underwent duodenal stent placement.⁵² Biliary stent dysfunction developed in 17 patients (52%), showing that this procedure was a risk factor for biliary stent dysfunction.

therapy, a setting in which stenting may be more risky.⁵⁰

Clinical outcomes

Stents can be placed successfully in over 90% of patients, with typical clinical success rates of 80% or higher.54-59 Technical failure is usually due to an inability to pass the guidewire through the stricture, anatomic difficulties such as severe looping within the dilated stomach, or complicated postsurgical anatomy.⁵⁰ Published experiences with enteral stenting for gastroduodenal obstructions consist mostly of observational studies, 54,57,58,60-69 although small randomized trials have compared stent placement with a surgical bypass for palliation of malignant gastric outlet obstruction.44,70,71 In one of these randomized trials, 39 patients with malignant gastric outlet obstruction were assigned to either gastrojejunostomy (18 patients) or enteral stent placement (21 patients).⁴⁴ Food intake improved more rapidly in the stent group compared to the surgery group (median, 5 days vs. 8 days). Nevertheless, long-term relief was worse in the stent group (median duration of relief, 50 days vs. 72 days). Major complications were more common in the stent group, though there were no differences between the groups in survival or health-related quality of life scores. In a second randomized trial, 18 patients were assigned to endoscopic stent placement or surgical bypass.⁷⁰ Endoscopic stenting was superior to surgery in the median operative time (40 minutes vs. 93 minutes), mean time to restoration of oral intake (2.1 days vs. 6.3 days), and median length of hospital stay (3.1 days vs. 10 days). There were no significant differences between the groups in morbidity, mortality, gastric emptying, or clinical outcomes at the 3-month follow-up. Observational studies suggest that enteral stenting has a success rate similar to that of surgical palliation, with approximately 90% of patients improving clinically, and is associated with lower morbidity, procedure-related mortality, and cost.^{55,56,72-74} A database study comparing 425 stenting procedures with 339 surgical bypasses for malignant gastric outlet obstruction found that the median length of stay was shorter for stenting procedures than for surgery (8 days vs. 16 days) and the median cost was lower.74 Furthermore, some studies suggest that stenting may permit a better quality of life compared to other forms of palliation (such as non-oral feeding through a jejunostomy tube).^{75,76} Studies of one of the newer stents (Evolution Duodenal Stent) have found similar success rates. In a series of 46 patients, technical success was achieved in 89% of cases, with a clinical success rate of 72%.⁶⁸

The following conclusions were obtained in a systematic review of 44 studies comparing stenting and gastrojejunostomy (1,046 patients receiving a stent, and 297 undergoing gastrojejunostomy).55 There were no significant differences between stent insertion and gastrojejunostomy in terms of technical success (96% vs. 100%), early complications (7% vs. 6%), late major complications (18% vs. 17%), or persistent symptoms (8% vs. 9%). Initial clinical success was higher after stent placement (89% vs. 72%), as were recurrent obstructive symptoms (18% vs. 1%). Mean survival was 105 days after stent placement and 164 days after gastrojejunostomy. Despite initial success, 15% to 40% of patients require reintervention for recurrent symptoms or biliary obstruction following stent placement. Reintervention rates were generally lower in patients undergoing gastrojejunostomy.44,55,69,73 Furthermore, some patients may not improve, even after successful stent placement. This may result from unidentified distal sites of malignant obstruction, diffuse peritoneal carcinomatosis with bowel encasement, or functional gastric outlet obstruction from neural (celiac axis) tumor involvement.^{15,50,77}

In a prospective observational study of 22 patients undergoing SEMS for benign pyloric stenosis, early symptom improvement was achieved in 18 patients (81.9%). However, during the follow-up period (mean 10.2 months), the stents remained in place in only seven patients (31.8%). Among the 15 patients with stent migration, seven patients (46.6%) showed continued symptom improvement without recurrent symptoms.⁷⁸ However, most publications in the benign disease have small sample sizes and are uncontrolled. The optimal indications, methods, and duration of stenting for benign diseases should to be determined in further studies.

CONCLUSIONS

Endoscopic stent placement in upper GI disorders is currently the most common modality for palliation of symptoms in patients with malignant upper GI obstruction, replacing conventional surgical bypass. Endoscopic stenting is associated with less invasiveness, high compliance, less complications, shorter hospital stays, and lower costs. Although there have been several complications, such as migration and recurrent obstruction, newly developed stents are expected to overcome these limitations and extend to a variety of benign and malignant disorders.

Conflicts of Interest _

The authors have no financial conflicts of interest.

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