

GUIDELINE



ASGE guideline on the role of endoscopy in the management of benign and malignant gastroduodenal obstruction



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This document was reviewed and approved by the Governing Board of the American Society for Gastrointestinal Endoscopy.

This American Society for Gastrointestinal Endoscopy guideline provides evidence-based recommendations for the endoscopic management of gastric outlet obstruction (GOO). We applied the Grading of Recommendations, Assessment, Development and Evaluation methodology to address key clinical questions. These include the comparison of (1) surgical gastrojejunostomy to the placement of self-expandable metallic stents (SEMS) for malignant GOO, (2) covered versus uncovered SEMS for malignant GOO, and (3) endoscopic and surgical interventions for the management of benign GOO. Recommendations provided in this document were founded on the certainty of the evidence, balance of benefits and harms, considerations of patient and caregiver preferences, resource utilization, and cost-effectiveness. (Gastrointest Endosc 2021;93:309-22.)

The clinical syndrome of gastric outlet obstruction (GOO) occurs as a result of a narrowing in the region of the gastroduodenum resulting in failed or delayed passage of gastric contents from stomach to jejunum. GOO typically presents with early satiety, weight loss, nausea, vomiting, and abdominal pain. Etiologies of GOO include both malignant and benign processes. Historically, peptic ulcer disease (PUD) was the most common cause of GOO. The development and use of acid-suppressing medication and the diagnosis and treatment of *Helicobacter pylori* has decreased the incidence of PUD, and thus malignancy is now the leading cause of GOO. ¹⁻⁴

The most common malignant causes of GOO are pancreatic, gastric, and duodenal cancer (Table 1). The worldwide incidence and mortality of gastric cancer is higher than pancreatic cancer.⁵ In the United States,

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however, pancreatic and gastric cancers are the 11th and 16th most common newly presenting cancers. Pancreatic cancer has a high mortality rate and is the third leading cause of cancer death in the United States. ^{6,7} GOO occurs in approximately 20% of cases of advanced pancreatic cancer, and it is estimated that more than 11,000 pancreatic cancer patients were diagnosed with GOO in 2019. ⁸⁻¹⁰ The most common benign etiology of GOO is PUD from either nonsteroidal anti-inflammatory drugs (NSAIDs) or *H pylori* infection. Other inflammatory, infectious, and iatrogenic causes of benign GOO are highlighted in Table 1.

AIMS AND SCOPE

The aim of this document is to provide evidence-based recommendations for the endoscopic management of GOO. The committee formulated clinical questions central to the endoscopic management of GOO, comparing clinical outcomes and adverse events with different treatment

TABLE 1. Etiology of gastric outlet obstruction						
Causes of benign gastric outlet obstruction	Causes of malignant gastric outlet obstruction					
Peptic ulcer disease	Pancreatic cancer					
Nonsteroidal anti-inflammatory drug-induced strictures	Gastric cancer					
Caustic ingestions	Duodenal cancer					
Chronic pancreatitis	Gastric carcinoids					
Pseudocysts	Small intestine carcinoids					
Walled-off pancreatic necrosis	Pancreas neuroendocrine tumors					
GI polyps	Lymphoma					
Lipomas	Sarcomas					
Surgical anastomotic strictures	GI stromal tumors					
Foreign bodies	Ampulla of Vater cancer					
Radiation-induced strictures	Metastatic cancer					
Eosinophilic enteritis	Cholangiocarcinoma					
Tuberculosis	Gallbladder cancer					

options. This document addresses the following clinical questions:

- 1. In patients with incurable malignant GOO undergoing a palliative intervention, what is the role of gastric and/or duodenal self-expandable metal stent (SEMS) placement compared with surgical gastrojejunostomy (GJ)?
- 2. In patients with incurable malignant GOO undergoing palliative endoscopic stent placement, what is the role of covered SEMS placement compared with uncovered SEMS placement?
- 3. In patients with benign GOO, what is the role of endoscopic management (balloon dilation and/or covered SEMS placement) compared with surgical GJ?

The panel also summarized the diagnosis and management considerations pertinent to mechanical GOO in the pediatric population.

METHODS

Overview

This guideline was conceptualized and created using the Grading of Recommendations, Assessment, Development and Evaluation (GRADE) framework, 11,12 beginning with formulation of clinical questions in the Population, Intervention, Comparison, and Outcome (PICO) format. For each PICO question, several outcomes were included and ranked according to their importance. Systematic reviews (SRs) of the available literature were performed for each clinical question. The quality or certainty in the evidence and strengths of recommendations were based on the GRADE framework. When existing SRs were identified, they were used to inform the guideline, when appropriate. If no existing SRs were found, a new SR and

meta-analysis (MA) was conducted, when possible, with the assistance of an expert librarian, and evidence profiles were then created with the guidance of a GRADE methodologist. Evidence profiles were presented to a multidisciplinary GRADE panel and members of the American Society for Gastrointestinal Endoscopy (ASGE) Standards of Practice committee. This document was approved by the ASGE Governing Board in May 2020.

Panel composition and conflict of interest management

The panel consisted of content experts (T.J., A.S.), a GRADE methodologist (B.Q.), a patient representative (D.B.), a GI oncologist (A.M.), a GI oncologic surgeon (M.T.), and members of the Standards of Practice committee. All panel members were required to disclose potential financial and intellectual conflicts of interest, which were addressed according to ASGE policies (https://www.asge.org/forms/conflict-of-interest-disclosure and https://www.asge.org/docs/default-source/about-asge/mission-and-governance/asge-conflict-of-interest-and-disclosure-policy.pdf). The panel meeting took place in Chicago on March 6, 2020.

Formulation of clinical questions

Three clinical questions were developed by the authors of the document and members of the ASGE Standards of Practice committee using the PICO format and approved by the ASGE Governing Board. For all clinical questions, potentially relevant patient-centered outcomes were identified and rated from not important to critical through a consensus process. Relevant clinical outcomes included tolerance of an oral diet, reintervention rate, duration of patency (reintervention-free period), major adverse event rate, SEMS migration rate, time to oral intake, and length of hospital stay. A list of the PICO questions is detailed in Table 2.

Literature search and study selection criteria

For each PICO question, a literature search for existing SRs and MAs was performed. Details of the search strategies are reported in Appendix 1 (available online at www. giejournal.org). An expert medical librarian (K.K.) performed all searches. Citations were imported into EndNote (Thompson Reuters, Philadelphia, Pa, USA), and duplicates were removed. The EndNote library was uploaded into Covidence (www.covidence.org), and 2 independent reviewers were assigned to each search. Each study was reviewed based on the title and abstract using explicit criteria for inclusion or exclusion in the GRADE analysis. As necessary, full texts were then reviewed and differences resolved by consensus. If existing SRs and MAs were available, inclusion and exclusion criteria were reviewed, and methodologic quality of the study was assessed using the AMSTAR 2 checklist (A MeaSurement Tool to Assess systematic Reviews; https://amstar.ca/

TABLE 2. Clinical questions addressed in the document using the Population, Intervention, Comparator, and Outcome (Population, Intervention, Comparator, and Outcomes) format

Population	Intervention	Comparator	Outcomes	Rating
1. Patients with incurable	Endoscopically placed SEMS	Surgical gastrojejunostomy	1. Survival	Critical
malignant GOO			2. Postoperative mortality	Critical
requiring palliation			3. Re-establishment of oral intake	Critical
			4. Time to oral intake re-establishment	Critical
			5. Length of hospital stay	Critical
			6. Patency	Critical
			7. Reintervention rate	Critical
			8. Adverse events	Critical
			9. Type of diet	Important
2. Patients with incurable	Covered SEMS	Uncovered SEMS	1. Tolerance of oral diet	Critical
malignant GOO requiring palliation by			2. Occlusion rate	Critical
endoscopically placed			3. Migration rate	Critical
SEMS			4. Reintervention rate	Critical
			5. Adverse events	Critical
3. Patients with benign	Endoscopic management (balloon	Surgery	1. Tolerance of oral diet	Critical
GOO undergoing intervention	dilation, covered SEMS)		2. Major adverse events	Critical
intervention			3. Reintervention rate	Critical

GOO, Gastric outlet obstruction; SEMS, self-expandable metal stent.

Amstar_Checklist.php).¹³ Only SRs and MAs meeting the quality thresholds were used for data synthesis. When applicable, available SRs and MAs were updated based on literature review as described above.

If no existing SR or MA was identified, a full SR and MA (when possible) was conducted using the recommendations of the Preferred Reporting Items for Systematic Reviews and Meta-analyses criteria. 14 A medical librarian performed a comprehensive literature search of Ovid Medline (Ovid MEDLINE in-process and other nonindexed citations, Ovid MEDLINE Daily, and Ovid MEDLINE 1946 to September 2018), Embase (via Embase.com 1947 to September 2018), and the Cochrane Database of Systematic Reviews/Cochrane Register of controlled trials (via Wiley Online Library). The searches were limited to human studies published in English and cited through December 2019. Citations were imported into EndNote (Thompson Reuters), and duplicates were removed. The EndNote library was then uploaded into Covidence. Two reviewers (T.J., A.S.) were assigned to each search for each PICO question. Studies were first screened by title and abstract and then by full text, and all conflicts were resolved by consensus. Abstracts and studies with fewer than 10 patients were excluded during screening.

Data extraction and statistical analysis

When necessary, data extraction was accomplished by using Microsoft Excel (Microsoft Corporation, Redmond, Wash, USA). The outcomes varied by PICO question as described

below. I² statistics were used to assess heterogeneity, and funnel plots were used to assess for publication bias.

Certainty in evidence (quality of evidence)

The certainty of the evidence was determined using the GRADE framework. The GRADE approach to rating the quality or certainty of evidence begins with the study design (randomized controlled trials [RCTs] or observational studies) and then reviews 5 parameters that may reduce quality rating (methodologic limitations, inconsistency, indirectness, imprecision, and publication bias) and 3 parameters that may elevate the rating (large effect, dose-response gradient, plausible confounding). The final quality of evidence scores range from very low to high (Table 3). Guideline developers then formulate the recommendations (guidance statements) and consider the direction (for or against) and grade the strength (strong or weak) of the recommendations based on the criteria outlined in the GRADE approach. Our GRADE evidence profiles, developed using GDTpro application (http://gdt.guidelinedevelopment.org/app), contain detailed information about the quality of evidence assessment and the summary of findings for each of the included outcomes.

Considerations in the development of recommendations

The main factors driving our recommendations included balance between benefits and harms while accounting for the best estimates of the magnitude of effects

TABLE 3. System for rating the quality of evidence

Quality of evidence	Definition	Interpretation
High	We are very confident that the true effect lies close to that of the estimate of effect.	Future research is very unlikely to change our confidence in the estimate of the effect.
Moderate	We are moderately confident in the effect estimate: The true effect is likely to be close to the estimate of effect, but there is a possibility that it is substantially different.	Further research is likely to have an impact on our confidence in the estimate of the effect and may change the estimate.
Low	Our confidence in the effect estimate is limited: The true effect may be substantially different from the estimate of effect.	Further research is very likely to have an impact on our confidence in the estimate of the effect and is likely to change the estimate.
Very low	We have very little confidence in the estimate of the effect: The true effect is likely to be substantially different from the estimate of effect.	Any estimate of the effect is very uncertain.

Adapted from Guyatt et al.11

and importance of outcomes, overall quality of evidence, confidence in values and preferences and their variability, and cost/resource implications. The final wording of the recommendations (including direction and strength), remarks, and qualifications were decided by consensus and were approved by all members of the panel. According to the GRADE approach, the recommendations are either "strong" or "conditional" (Table 4). The words "the guideline panel recommends" are used for strong recommendations and "suggests" for conditional recommendations. ¹⁵

RESULTS

Question 1: In patients with incurable malignant GOO undergoing a palliative intervention, what is the role of gastric and/or duodenal SEMS placement compared with surgical GJ?

Recommendation: In patients with incurable malignant GOO undergoing palliative intervention, we suggest either SEMS placement or surgical GJ. The selected approach should be based on patient characteristics, preferences, multidisciplinary input, and local expertise. (Conditional recommendation, low quality of evidence)

Comment: Based on shared decision-making, in patients who are poor surgical candidates with short life expectancy (<6 months) and those who place a high value on resumption of oral diet and being discharged early, we suggest SEMS placement compared with surgical GI. In patients with a life expectancy of >6 months and good performance status, we suggest surgical GI compared with SEMS placement.

Summary of the evidence

Our literature search identified 9 SRs, ¹⁶⁻²⁴ of which 3 were rated as moderate quality based on AMSTAR 2 criteria. ^{20,21,23} Two reviews ^{20,23} exclusively included the 3 published RCTs comparing surgical GJ with SEMS placement for malignant GOO. ²⁵⁻²⁷ Of the 3 included

RCTs with a total of 84 patients, 2 were single-center trials^{25,27} and 1 was a multicenter trial.²⁶ The panel decided to use 2 SRs^{21,23} to guide this PICO question. The study by Upchurch et al²³ is a Cochrane Library SR and MA that included outcomes aligned with our outcomes of interest, as decided a priori. The second SR and MA, by Mintziras et al,²¹ included 24 retrospective, comparative studies in addition to the 3 RCTs, for a total of 2354 patients. The Risk Of Bias In Non-randomized Studies of Interventions tool²⁸ for assessing risk of bias was used in this study (Table 5).

For the outcome of survival, only 1 RCT provided comparative data, which was reported as median number of days for SEMS placement versus surgical GJ.²⁶ There was no statistically significant difference in survival between SEMS versus surgical GJ (mean difference [MD], 22 days shorter; 95% confidence interval [CI], 53.45 shorter to 9.45 days more). However, review of 9 observational studies reported a survival benefit with surgical GJ (MD, 42.85 days; 95% CI, 12-73.7).²¹ The outcome of in-hospital mortality was reported in 1 of 3 RCTs, which reported no difference between the 2 groups (risk ratio [RR], 0.72; 95% CI, 0.14-3.64).²⁷ Similarly, the SR by Mintziras et al²¹ reviewed 15 studies and reported no difference in procedure-related mortality (odds ratio, 0.55; 95% CI, 0.27-1.16).

For the outcome of re-establishment of oral intake, there was no difference between SEMS placement and surgical GJ (RR, 0.98; 95% CI, 0.88-1.09).²³ However, the time to resumption of oral intake was sooner in those undergoing SEMS placement compared with surgery based on 2 RCTs (MD, 3.07 fewer days; 95% CI, 4.76-1.39).^{25,26} Similarly, 11 observational studies that included 826 patients demonstrated that patients who underwent endoscopic SEMS placement resumed oral intake nearly 5 days earlier than those who had surgical GJ (MD, 4.9 days fewer; 95% CI, 6.75-3.05).²¹

The 3 RCTs showed a significantly shorter hospital length of stay with SEMS placement versus surgical GJ (MD, 6.7 days shorter; 95% CI, 9.41-3.98). Similar results were noted

TABLE 4. Interpretation of definitions of strength of recommendation using Grading of Recommendations, Assessment, Development and Evaluation framework

Implications for	Strong recommendation	Conditional recommendation
Patients	Most individuals in this situation would want the recommended course of action, and only a small proportion would not.	Most individuals in this situation would want the suggested course of action, but many would not.
Clinicians	Most individuals should receive the intervention. Formal decision aids are not likely to be needed to help individual patients make decisions consistent with their values and preferences.	Recognize that different choices will be appropriate for individual patients and that you must help each patient arrive at a management decision consistent with his or her values and preferences. Decision aids may be useful in helping individuals to make decisions consistent with their values and preferences.
Policymakers	The recommendation can be adopted as policy in most situations. Compliance with this recommendation according to the guideline could be used as a quality criterion or performance indicator.	Policymaking will require substantial debate and involvement of various stakeholders.

Adapted from Andrews et al.¹⁵

when observational studies were included (MD, 9.75 days shorter; 95% CI, 11.6-7.9 days).²¹ When assessing duration of patency, defined as time to recurrence of obstructive symptoms, there was no statistically significant difference between the 2 groups (RR, 5.08; 95% CI, 0.96-26.74).^{25,26} However, reintervention rates were higher in SEMS placement versus surgical GJ (odds ratio, 2.95; 95% CI, 1.70-5.14) based on 14 observational studies and 1 RCT.²¹ Based on 3 RCTs, there was no significant difference in serious adverse events (Clavien-Dindo classification grade III and above)²⁹ between the 2 groups for SEMS placement versus surgical GJ (RR, 1.15; 95% CI, 0.33-3.98).²³ Similar results were noted in an analysis that included observational studies (RR, 0.73; 95% CI, 0.50-1.06).²¹ The risk of adverse events requiring need for intervention (including endoscopy) was significantly higher with SEMS placement compared with the surgical GJ group (RR, 4.71; 95% CI, 1.36-16.3).²³

Regarding diet, 2 of 3 RCTS reported pre- and postintervention scores on the Gastric Outlet Obstruction Scoring System (GOOSS). Fiori et al 25 reported that 8 of 9 patients were able to tolerate a soft diet 2 days after SEMS placement and a regular diet 3 days afterward. In comparison, 6 of 9 surgical patients tolerated a soft diet on the sixth postoperative day and a regular diet on the seventh postoperative day. Jeurnink et al 26 reported outcomes as either a GOOSS score \geq 2 (soft solids to regular diet) or not for both groups. Seven retrospective comparative studies included GOOSS scores, but our review of outcomes did not show a significant difference in type of oral diet tolerated for SEMS placement versus surgery. $^{30-36}$

Certainty in the evidence

The overall certainty for all outcomes results reported by the RCTs was moderate. These outcomes included survival, postoperative mortality, re-establishment of oral intake, time to oral intake resumption, length of hospital stay, patency, reintervention rate, serious adverse events, and serious adverse events with need for reintervention. We downgraded for imprecision because of the small number of events in the outcomes. The overall certainty of the observational review²¹ was very low; downgraded for risk of bias and inconsistency as detailed in Table 5.

Considerations

Our literature search did not identify studies citing patient preferences for SEMS placement versus surgical GJ before palliation of unresectable malignant GOO. Patient surveys of quality of life and satisfaction were performed after intervention in each of the 3 RCTs, 2 of which showed no difference in patient satisfaction after SEMS placement versus surgical GJ. 25,26 Mehta et al 27 reported no significant decrease in pain or mental health score 1 month after either intervention; however, there was a significant improvement in the mean physical health score (as measured by the 36-item Short-Form Health Survev)³⁷ 1 month after SEMS placement compared with after surgery. Models reviewing prognostic factors for survival and performance scores suggest that SEMS placement was preferred over surgical GJ in those with shorter life expectancies.34,38,39

The panel considered the evidence on cost-effectiveness comparing the 2 treatment options. Most studies compared initial interventional costs but did not account for total costs after long-term follow-up. Initial costs for SEMS placement were less than for surgical GJ, primarily because of shorter initial hospital length of stay. A decision analysis model comparing cost-effectiveness of enteral SEMS versus surgical GJ concluded that endoscopic stent placement (\$8213) was less costly than open (\$12,191) and laparoscopic GJ (\$10,340) over a 1-month period of follow-up. The success rate and mortality rates used with the model favored SEMS placement over surgery. A retrospective review of a claims database for employer-provided insurance and Medicare supplemental insurance

		Cei	rtainty assess	ment			No. of	patients		Effect		
	Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other consid- erations	SEMS	Surgical GJ	Relative (95% CI)	Absolute (95% CI)	Certainty	Importanc
Surviva 1	Randomized trial [Jeurnick]	Not serious	Not serious	Not serious	Serious*	None	21	18	-	MD 22 days lower (53.45 lower to 9.45 higher)	⊕⊕⊕⊝ MODERATE	CRITICAL
Surviva	al									9,		
9	Observational studies	Serious†	Serious‡	Not serious	Not serious	None	297	216	-	MD 42.85 days fewer (73.7 fewer to 12 fewer)	⊕○○○ VERY LOW	CRITICAL
Postop	erative mortali	ty										
1	Randomized trial [Mehta]	Not serious	Not serious	Not serious	Serious*	None	2/13 (15.4%)	3/14 (21.4%)	RR 0.72 (0.14 to 3.64)	50 fewer per 1,000 (from 178 fewer to 284 more)	⊕⊕⊕⊝ MODERATE	CRITICAL
Postop	erative mortali	ty										
15	14 Observational studies 1 Randomized trial [Mehta]	Serious†	Not serious	Not serious	Not serious	None	27/443 (6.1%)	44/515 (8.5%)		37 fewer per 1,000 (from 61 fewer to 12 more)	⊕○○○ VERY LOW	CRITICAL
Re-esta	blishment of o	ral intak	e									
3	Randomized trials [Fiori, Mehta,Jeurnick]	Not serious	Not serious	Not serious	Serious*	None	39/42 (92.9%)	39/40 (97.5%)	RR 0.98 (0.88 to 1.09)	20 fewer per 1,000 (from 117 fewer to 88 more)	⊕⊕⊕○ MODERATE	CRITICAL
Time to	oral intake re	sumptio	n									
2	Randomized trials [Fiori, Jeurnick]	Not serious	Not serious	Not serious	Serious*	None	30	27	-	MD 3.07 days fewer (4.76 fewer to 1.39 fewer)	⊕⊕⊕⊝ MODERATE	CRITICAL
Time to	o oral intake re	sumptio	n									
11	Observational studies	Serious†	Serious‡	Not serious	Not serious	None	544	282	-	MD 4.9 days fewer (6.75 fewer to 3.05 fewer)	⊕○○○ VERY LOW	CRITICAL
Length	of hospital sta	ıy										
3	Randomized trials [Fiori, Jeurnick, Mehta]	Not serious	Not serious	Not serious	Serious*	None	41	43	-	MD 6.7 days fewer (9.41 fewer to 3.98 fewer)	⊕⊕⊕○ MODERATE	CRITICAL
Length	of hospital sta	ıy										
20	19 Observational studies 1 Randomized trial [Mehta]	Serious†	Serious‡	Not serious	Not serious	None	882	689	-	MD 9.75 days fewer (11.6 fewer to 7.9 fewer)	⊕○○○ VERY LOW	CRITICAL
Patenc	у											
2	Randomized trials [Fiori, Jeurnick]	Not serious	Not serious	Not serious	Serious*	None	8/30 (26.7%)	1/27 (3.7%)	RR 5.08 (0.96 to 26.74)	151 more per 1,000 (from 1 fewer to 953 more)	⊕⊕⊕○ MODERATE	CRITICAL

TABLE 5. Continued

		Cer	tainty assessi	ment			No. of	patients		Effect		
	s Study design	Risk of bias	Inconsistency	Indirectness	Imprecision	Other consid- erations	SEMS	Surgical GJ	Relative (95% CI)	Absolute (95% CI)	Certainty	Importance
Reinte	rvention rate											
15	14 Observational studies 1 Randomized trial [Jeurnick]	Serious†	Serious‡	Not serious	Not serious	None	250/ 955 (26.2%)	84/749 (11.2%)	OR 2.95 (1.70 to 5.14)	159 more per 1,000 (from 65 more to 282 more)	⊕○○○ VERY LOW	CRITICAL
Advers	se events, serio	us (Majoı	complication	ıs)								
3	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	6/43 (14.0%)	4/41 (9.8%)	RR 1.15 (0.33 to 3.98)	15 more per 1,000 (from 65 fewer to 291 more)	⊕⊕⊕⊝ MODERATE	CRITICAL
Advers	se events, serio	us (Majoı	complication	s)								
21	19 Observational studies 2 Randomized trials [Fiori, Jeurnink]	Serious†	Not serious	Not serious	Not serious	None	120/ 1099 (10.9%)	113/902 (12.5%)	OR 0.73 (0.50 to 1.06)	31 fewer per 1,000 (from 58 fewer to 7 more)	⊕○○○ VERY LOW	CRITICAL
Advers	se events, serio	us (Majoı	complication	s): with nee	d for reinte	rvention						
3	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	13/43 (30.2%)	2/41 (4.9%)	RR 4.71 (1.36 to 16.30)	181 more per 1,000 (from 18 more to 746 more)	⊕⊕⊕○ MODERATE	CRITICAL

CI, Confidence interval; MD, mean difference; OR, odds ratio; RR, risk ratio.

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plans identified patients who received palliative care for malignant GOO from 2012 to 2015. 41 The median 90-day cost of SEMS placement and associated readmissions was \$18,500 (range, \$4100-\$49,500) compared with \$37,200 (range, \$12,200-\$67,800) for surgery. In the only cost analysis conducted together with an RCT, initial costs were higher for surgical GJ (\$9782) than for SEMS placement (\$5035) because of the costs associated with longer hospital stay. 42 Comparison of follow-up costs found no significant difference between the 2 groups. However, longterm relief of obstruction was better with surgical GJ, with more patients requiring repeat interventions for obstructive symptoms after SEMS placement. This study suggested that surgical GJ might be beneficial over SEMS placement for patients with life expectancy of ≥ 2 months, with a calculated incremental cost-effectiveness ratio of \$192 per day with a GOOSS score ≥ 2 (soft solids to regular diet) adjusted for survival.

Discussion

The panel reviewed and discussed the quality of evidence and weighed the benefits and potential harms. It should be highlighted that this clinical question only addresses the application of SEMS compared with surgical

GJ in the palliation of unresectable cancer and not as a bridge to surgery. With regard to the outcomes assessed, there was no difference in success rates, major adverse event rates, and postoperative mortality between the 2 interventions. Clinically meaningful differences of shorter time to resumption of oral intake and shorter length of hospital stay were reported with SEMS. These outcomes may be helpful in discussion with patients and other healthcare providers when both surgical GJ and SEMS placement are available options. In addition, shorter time to resumption of oral intake and shorter recovery time with SEMS placement may decrease the time interval before palliative chemotherapy can be administered, 43,44 because maturation of the surgical GJ typically requires several days. This clinical question was not designed to compare outcomes between laparoscopic and open surgical GI.

It should be noted that reintervention rates for SEMS placement were significantly higher than for surgical GJ. The panel therefore agreed that although stent placement may offer a short-term advantage, patients whose performance status is good and whose life expectancy is longer than 6 months may benefit more from surgical GJ than SEMS placement. Because of the complexity of this

^{*}small number of events

[†]Bias due to confounding and selection of participants

decision-making, the panel recommended that multidisciplinary evaluation of a patient's performance status, projected clinical course, and preferences should guide the decision to palliate by SEMS placement versus surgical GJ. The practice of performing surgical GJ in those with longer life expectancy has not been validated. However, it is a common clinical approach and may be a source of bias in observational studies that report longer survival times in patients undergoing surgical GJ compared with SEMS placement.²¹ Similarly, a recent, large, comparative, retrospective, single-center study reported longer survival with surgical GJ than SEMS placement, with poor performance status, ascites, and low albumin being predictors of failure. 45 However, comparison of the 2 groups suggested that patients who underwent surgery had higher albumin and performance status, and like other retrospective studies, selection bias may have affected the outcome favoring surgery.

Although it was not included in the initial PICO question, the panel acknowledged the evolving role of EUS-guided gastroenterostomy (EUS-GE) for the management of malignant GOO. EUS-GE is a relatively new technique that is increasingly used as a therapeutic option for palliation of malignant GOO at high-volume tertiary care centers. EUS is used to identify and access the duodenum or jejunum beyond the area of obstruction from within the stomach, and placement of a lumenapposing metal stent (LAMS) is performed to create a GE. At the time of drafting this guideline document, the application of LAMS for this purpose was considered an off-label use.

A SR and MA of 5 studies with 199 patients who underwent EUS-GE for both benign and malignant GOO reported a technical success rate of 92.6% (95% CI, 88.26-95.79; $I^2 = 0\%$); serious adverse event rate of 5.61% (95% CI, 2.87-10.67; $I^2 = 1.67\%$), including occurrences of stent maldeployment, peritonitis, surgical intervention, and hemoperitoneum; and reintervention of 11.43% (95% CI, 7.29%-17.46%; $I^2 =$ 17.38%).⁴⁷ single-center, Α retrospective comparing EUS-GE (n = 22) with endoscopic gastroduodenal SEMs (n = 78) for palliation of malignant GOO reported 100% technical success in both groups and similar length of hospital stay but a higher reintervention rate in the SEMS group (32% vs 8.3%, P =0.021).⁴⁸ Follow-up times for both groups were limited because the authors elected to perform empiric exchange of the LAMS for those patients still alive at 6 months. Another retrospective study comparing EUS-GE performed at multiple centers with SEMS placement at 1 center for malignant GOO found similar results. 49 A retrospective study compared EUS-GE (n = 93) with surgical GJ (n = 63) for palliation of malignant GOO⁵⁰ and found the technical success rate of surgical GJ was superior (100% vs 87%, P = 0.009) with no difference in length of hospital stay, serious

adverse event rates, rates of recurrent GOO, and mean time to reintervention between the 2 groups.

Relative contraindications to EUS-GE include involvement of the intended transgastric access site with tumor or blood vessels including perigastric varices, large volume ascites, peritoneal carcinomatosis, and bowel obstruction distal to the intended site of GE placement. Prospective RCTs are needed to compare EUS-GE with both surgical GJ and endoscopic SEMS placement.

Question 2: In patients with incurable malignant GOO undergoing palliative endoscopic stent placement, what is the role of covered SEMS placement as compared with uncovered SEMS placement?

Recommendation: In patients with incurable malignant GOO undergoing palliative endoscopic stent placement, there is insufficient evidence to preferentially recommend covered over uncovered SEMS. The final decision should be based on regional stent availability, patient characteristics, and patient preferences. (Conditional recommendation, moderate quality of evidence)

Summary of the evidence

Our literature search identified 4 SRs of which 3 were rated as critically low in quality based on AMSTAR 2 criteria. ^{20,51-53} The SR with MA conducted by Minata et al²⁰ was moderate in quality and included 5 RCTs comparing covered with uncovered SEMS. One study was single center, ⁵⁴ 1 was conducted at 2 centers, ⁵⁵ and the remaining were multicenter studies. ⁵⁶⁻⁵⁸ Three of the 5 studies only included patients with gastric cancer, ^{54,56,58} accounting for 247 of 443 cases (55.8%). All 5 studies were conducted in East Asian countries, because covered enteral stents for gastroduodenal obstruction are not available for clinical use in the United States. The evidence profile for this PICO is summarized in Table 6.

Based on 5 RCTs, there was no difference in successful resumption of an oral diet between covered (204/221; 92.3%) and uncovered (201/222; 90.5%) SEMS (risk difference [RD], 0.02; 95% CI, -0.03 to 0.07). However, the RCTs also showed that SEMS occlusion occurred at a significantly lower rate with covered (9/221; 4.1%) compared with uncovered (56/222; 25.2%) SEMS (RD, -0.21; 95% CI, -0.27 to -0.15). This means that the rate of occlusion in covered SEMS was far lower than for uncovered SEMS. In contrast, SEMS migration occurred at a significantly higher rate with covered (26/221; 11.8%) compared with uncovered (6/222; 2.7%) SEMS (RD, 0.09; 95% CI, 0.04 to 0.14).

Reintervention rates were recorded in only 3 studies, ⁵⁶⁻⁵⁸ and no difference was reported between the 2 groups (covered, 26/150 [17.3%]; uncovered, 30/151 [19.9%]; RD, -0.03; 95% CI, -0.11 to 0.06). Covered SEMS had an adverse event rate of 22.3% (42/188) and uncovered SEMS a rate of 30.5% (58/190), with an RD of -0.08 (95% CI, -0.17 to 0.0). One study reported a disproportionately high adverse event rate in the covered SEMS

TABLE 6. Evidence profile for covered versus uncovered SEMS for palliation of incurable malignant GOO

		Certai	nty assessme	nt		No. of pat	o. of patients Effect				
No. of studies	Study design	Risk of bias		Indirectness	Imprecision co	Other nsiderations	Covered SEMS	Uncovered SEMS	Relative (95% CI)	Certainty	Importance
Toleran	nce of oral die	t									
5	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	204/221 (92.3%)	201/222 (90.5%)	RD 0.02 (-0.03 to 0.07)	⊕⊕⊕⊝ MODERATE	CRITICAL
SEMS o	occlusion										
5	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	9/221 (4.1%)	56/222 (25.2%)	RD -0.21 (-0.27 to -0.15)	⊕⊕⊕⊝ MODERATE	CRITICAL
SEMS n	nigration										
5	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	26/221 (11.8%)	6/222 (2.7%)	RD 0.09 (0.04 to 0.14)	⊕⊕⊕⊜ MODERATE	CRITICAL
Reinter	vention rate										
3	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	26/150 (17.3%)	30/151 (19.9%)	RD -0.03 (-0.11 to 0.06)	⊕⊕⊕○ MODERATE	CRITICAL
Adverse	e events									<u> </u>	
5	Randomized trials	Not serious	Not serious	Not serious	Serious*	None	42/188 (22.3%)	58/190 (30.5%)	RD -0.08 (-0.17 to 0.00)	⊕⊕⊕⊝ MODERATE	CRITICAL

CI, Confidence interval

group of 85% (28/33) versus 34% (11/32) for uncovered SEMS.⁵⁸ Abdominal pain accounted for most events (13 covered vs 1 uncovered), followed by bleeding (11 covered vs 2 uncovered). In this study, the covered SEMS were tailored with a large funnel-shaped antimigration flange on the proximal (upstream) ends—which the uncovered SEMS did not feature. Whether this stent design contributed to the higher rate of abdominal pain or bleeding is unclear. The other 4 studies in the analysis compared similarly shaped SEMS. Because of its outlying results, the study with the uniquely tailored covered stents⁵⁸ was not included in the adverse event rate comparisons. After exclusion, no statistically significant difference in adverse event rates was noted between covered versus uncovered SEMS.

Certainty in the evidence

The overall certainty for all outcomes results reported by these RCTs was moderate. We downgraded for imprecision because of the small number of events. Although a risk of bias may occur with the inability to blind the study team to the type of SEMS placed, we decided not to downgrade on that basis because the studies had no major methodologic issues. Another potential source of bias was that 3 RCTs only included patients with gastric cancer, and 1 study had over 80%⁵⁷ affected by gastric cancer. Therefore, the application of these data to other etiologies of malignant GOO is unclear. In addition, although the adverse event rates were not significantly different between covered and uncovered SEMS, the severity of adverse events was not accounted for in the MA.

Considerations

Available literature demonstrates that both covered and uncovered SEMS had similar success and adverse event rates, with the primary differences in performance being occlusion of uncovered SEMS and migration of covered SEMS. In addition, the panel decided to compare severe adverse event rates between the 2 groups. The selected SR and MA that only included RCTs did not report a significant difference in rates of perforation (covered, 2/221 [0.9%]; uncovered, 0/222 [0%]; RD, 0.01; 95% CI, -0.01 to 0.03).²⁰ To assess the differences in adverse events, we performed a new SR that included the 5 RCTs and 8 additional observational studies⁵⁹⁻⁶⁶ with a total of 709 patients who underwent covered and 874 uncovered SEMS placement (Supplementary Table 1, available online at www. giejournal.org). All studies were conducted in East Asia, and most patients underwent SEMS placement for palliation of unresectable gastric cancer, with unresectable pancreatic cancer the second most common malignancy treated. Bleeding rates were no different between uncovered and covered SEMS when excluding for the study with the outlier funnel-shaped covered SEMS.⁵⁸ Similarly, there was no difference in rate of pancreatitis: 4 episodes of pancreatitis occurred, with 2 events in each group (covered, 2/155 [1.3%]; uncovered, 2/157 [1.3%]). However, only 2 studies evaluated for pancreatitis as an outcome. 60,64 Only 1 study evaluated for cholangitis as an adverse event, with only 2 events, 1 in each group (covered, 1/126 [0.8%]; uncovered, 1/126 [0.8%]). 60 Although perforations occurred at a higher rate with covered SEMS (10/448, 2.2%) than with uncovered SEMS

^{*}small number of events

(3/496, 0.6%), the studies did not detail whether the perforations occurred in the setting of migration.

No studies comparing costs between covered versus uncovered SEMS were identified. A covered enteral stent designed (and U.S. Food and Drug Administration approved) for palliation of malignant GOO is not available in the United States at this time; thus, all known comparative studies were conducted in East Asia.

Discussion

No difference in technical and clinical success, reintervention rates, or adverse event rates between covered versus uncovered SEMS are evident, but whether there is a significant difference in severe adverse events remains unclear based on our SR of available literature—a critical outcome for this PICO question. Occlusion, which occurred more often in uncovered SEMS, may not be as difficult to address endoscopically, because endoscopic ablation and placement of another SEMS within the occluded SEMS has been described.⁶⁷ Conversely, migration, which occurred more often with covered SEMS, may be more difficult to treat and may carry a higher risk for perforation.⁶⁸ Repositioning or removing migrated SEMS has been described, with placement of a new SEMS when repositioning fails.⁶⁹ The role of anchoring of covered duodenal SEMS by clips or endoscopic suturing to prevent migration has yet to be of proven benefit. 70,7 Another limitation for the reported outcomes was the heterogeneity in the different types of covered and uncovered SEMS included in the SR. These uncertainties in the available literature played a significant role in the panel's decision to not recommend one type of stent over the other. Future large, adequately powered RCTs comparing covered and uncovered SEMS in the United States are needed.

Question 3: In patients with benign GOO, what is the role of endoscopic management compared with surgical management?

Recommendation: There is insufficient evidence to support endoscopic management over surgical management of benign GOO. (Conditional recommendation, low quality of evidence)

Comment: Factors that should be considered in determining appropriate management include the etiology of benign GOO, length of the stricture, response to endoscopic balloon dilation (EBD), and health and comorbidities of the patients.

Summary of the evidence

Our literature search did not identify comparative studies between endoscopic versus surgical management of benign GOO. Rather, endoscopic management is typically attempted first, with the goal of successfully resolving benign GOO, whereas surgery is reserved for "salvage" therapy if endoscopic interventions prove insufficient or if a severe adverse event after endoscopic therapy occurs

that requires surgical intervention (ie, perforation). The approach to surgical interventions for the treatment of benign GOO may vary, depending on etiology of the stricture and available expertise. With the introduction of acid suppression therapy and recognition of $H\ pylori$ as a cause of PUD, surgery for benign GOO is uncommon. In the modern era, surgical pyloroplasty and laparoscopic GJ have been reported as acceptable treatments with low morbidity and mortality. $^{72-78}$

We performed an SR of EBD of benign strictures causing GOO. We identified 14 retrospective studies that included 596 patients.^{3,79-91} (Supplementary Table 2, available www.giejournal.org). at The significant heterogeneity in reported studies precluded pooling of data to achieve an estimate of effect. However, the results suggest that patients with benign GOO because of PUD have a high rate of response to endoscopic management without the need for surgery when inflammatory factors such as smoking and NSAIDs are discontinued and H pylori infection is treated. Corrosive strictures and strictures caused by chronic pancreatitis appear to be less responsive to endoscopic therapy. Strictures that require multiple dilations were also less likely to respond to endoscopic therapy.86

We also identified 2 studies describing the use of covered SEMS for benign GOO strictures. A series from Korea with 22 patients who underwent covered SEMS placement for benign GOO (21/22 from PUD) reported an 81.8% rate of early improvement in GOOSS, which was sustained in 67% of those patients. However, a high rate of SEMS migration was reported (62.5%, n=15). The use of covered SEMS in the United States for benign disease is off-label. All dedicated enteral stents in the United States are uncovered and should not be used for benign disease because they are not removable. Successful management of short, benign peptic strictures by off-label use of covered LAMS has been described in numerous case series and within SRs of LAMS used for therapy of GI tract strictures.

Discussion

Given the significant heterogeneity in surgical and endoscopic management of benign GOO, an indirect comparative analysis between these 2 approaches was not feasible. The panel acknowledges the need to assess for and address factors associated with response rates to EBD (*H pylori* infection, use of NSAIDs, and smoking), as highlighted in Supplementary Table 2. Patients with benign GOO secondary to long strictures (>3 cm), caustic ingestion, and chronic pancreatitis are less likely to respond to EBD. The need for multiple EBD procedures is predictive for treatment failure and need for surgery. Future prospective trials are needed to compare endoscopic and surgical approaches for management of patients with benign GOO. Similarly, comparative studies between EBD versus covered SEMS

(including the placement of LAMS) for benign GOO are required.

CONSIDERATIONS FOR THE PEDIATRIC POPULATION

GOO in pediatric patients is usually from benign etiologies. The most common cause is infantile hypertrophic pyloric stenosis, typically presenting with nonbilious emesis. Standard endoscopy is only used to exclude other diagnoses outside of infancy, and EUS has been used in some cases. Surgery is considered to be the standard management for infantile hypertrophic pyloric stenosis. Beyond infancy, peptic and caustic ulceration are the most common causes of GOO. Anatomic malformations including antral webs and duplication cysts as well as bezoars have been reported, and GOO as a manifestation of malignancy is rare in children. EBD has been used for therapy in pediatric GOO, and there are limited reports of needle-knife incisional therapy and SEMS placement. 101-105

SUMMARY AND CONCLUSIONS

Mechanical GOO occurs frequently with advanced malignancies and commonly affects those with progressive pancreatic and gastric cancer. SEMS placement and surgical GJ are acceptable treatment options, and multidisciplinary care, patient characteristics and preferences, and available local expertise should guide current treatment. Further investigation of outcomes from covered SEMS and EUS-GE would likely guide future management of malignant GOO. Ideal management of benign GOO remains uncertain, because prior studies focused on endoscopic management to avoid surgery rather than prospective comparison. Retrospective studies suggest higher rates of success with EBD when *H pylori* is treated and when NSAIDs and smoking are stopped and lower success rates with corrosive strictures and strictures requiring multiple dilations.

RECOMMENDATIONS

1. In patients with incurable malignant GOO undergoing palliative intervention, we suggest either SEMS placement or surgical GJ. The selected approach should be based on patient characteristics and preferences, multi-disciplinary input, and local expertise. (Conditional recommendation, low quality of evidence)

Comment: Based on shared decision-making, in patients who are poor surgical candidates with short life expectancy (<6 months) and those who place a high value on resumption of oral diet and being discharged early, we suggest SEMS placement compared with surgical GJ. In patients with a life expectancy of >6 months and good performance status, we suggest surgical GJ compared with SEMS placement.

- In patients with incurable malignant GOO undergoing palliative endoscopic stent placement there is insufficient evidence to preferentially recommend either covered or uncovered SEMS. The final decision should be based on regional stent availability, patient characteristics, and patient preference. (Conditional recommendation, moderate quality of evidence)
- There is insufficient evidence to uniformly support either endoscopic or surgical management for mixed varieties of benign GOO. (Conditional recommendation, low quality of evidence)

Comment: Factors that should be considered in determining appropriate management include etiology of benign GOO, length of stricture, and response to initial EBD.

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Abbreviations: AMSTAR, A MeaSurement Tool to Assess systematic Reviews, ASGE, American Society for Gastrointestinal Endoscopy; Cl, confidence interval; EBD, endoscopic balloon dilation; GE, gastroenterostomy; GJ, gastrojejunostomy; GOO, gastric outlet obstruction; GOOSS, Gastric Outlet Obstruction Scoring System; GRADE, Grading of Recommendations, Assessment, Development and Evaluation; IAMS, lumen-apposing metal stent; MD, mean difference; MA, meta-analysis; NSAID, nonsteroidal anti-inflammatory drug; PICO, Population, Intervention, Comparator, and Outcome; PUD, peptic ulcer disease; RCT, randomized controlled trial; RD, risk difference; RR, risk ratio; SEMS, self-expandable metal stent; SR, systematic review.

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REFERENCES

- Awan A, Johnston DE, Jamal MM. Gastric outlet obstruction with benign endoscopic biopsy should be further explored for malignancy. Gastrointest Endosc 1998;48:497-500.
- Feinstein LB, Holman RC, Yorita Christensen KL, et al. Trends in hospitalizations for peptic ulcer disease, United States, 1998-2005. Emerg Infect Dis 2010;16:1410-8.
- Misra SP, Dwivedi M, Misra V. Malignancy is the most common cause of gastric outlet obstruction even in a developing country. Endoscopy 1998;30:484-6.
- Shone DN, Nikoomanesh P, Smith-Meek MM, et al. Malignancy is the most common cause of gastric outlet obstruction in the era of H2 blockers. Am J Gastroenterol 1995;90:1769-70.
- Bray F, Ferlay J, Soerjomataram I, et al. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. CA Cancer J Clin 2018;68:394-424.
- National Cancer Institute. Estimated new cancer cases and deaths for 2018. SEER (Surveillance Epidemiology and End Results) cancer statistics review. Available from: http://www.seer.cancer.gov.
- American Cancer Society. Cancer Statistics Center. Available at https://cancerstatisticscenter.cancer.org. Accessed July, 2019.
- Manuel-Vazquez A, Latorre-Fragua R, Ramiro-Perez C, et al. Laparoscopic gastrojejunostomy for gastric outlet obstruction in patients with unresectable hepatopancreatobiliary cancers: a personal series and systematic review of the literature. World J Gastroenterol 2018;24:1978-88.
- 9. Sarr MG, Cameron JL. Surgical management of unresectable carcinoma of the pancreas. Surgery 1982;91:123-33.
- Singh SM, Longmire WP Jr, Reber HA. Surgical palliation for pancreatic cancer. The UCLA experience. Ann Surg 1990;212:132-9.
- Guyatt G, Oxman AD, Akl EA, et al. GRADE guidelines: 1. Introduction-GRADE evidence profiles and summary of findings tables. J Clin Epidemiol 2011;64:383-94.
- 12. Wani S, Sultan S, Qumseya B, et al. The ASGE'S vision for developing clinical practice guidelines: the path forward. Gastrointest Endosc 2018;87:932-3.
- Shea BJ, Reeves BC, Wells G, et al. AMSTAR 2: a critical appraisal tool for systematic reviews that include randomised or non-randomised studies of healthcare interventions, or both. BMJ 2017;358:j4008.
- 14. Moher D, Liberati A, Tetzlaff J, et al. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. Ann Intern Med 2009;151:264-9; W264.
- Andrews JC, Schunemann HJ, Oxman AD, et al. GRADE guidelines: 15.
 Going from evidence to recommendation-determinants of a recommendation's direction and strength. J Clin Epidemiol 2013;66: 736-25
- 16. Bian SB, Shen WS, Xi HQ, et al. Palliative therapy for gastric outlet obstruction caused by unresectable gastric cancer: a meta-analysis comparison of gastrojejunostomy with endoscopic stenting. Chin Med J (Engl) 2016;129:1113-21.
- Hosono S, Ohtani H, Arimoto Y, et al. Endoscopic stenting versus surgical gastroenterostomy for palliation of malignant gastroduodenal obstruction: a meta-analysis. J Gastroenterol 2007;42: 283-90.
- Jeurnink SM, van Eijck CH, Steyerberg EW, et al. Stent versus gastrojejunostomy for the palliation of gastric outlet obstruction: a systematic review. BMC Gastroenterol 2007;7:18.

- Ly J, O'Grady G, Mittal A, et al. A systematic review of methods to palliate malignant gastric outlet obstruction. Surg Endosc 2010;24: 290-7.
- Minata MK, Bernardo WM, Rocha RS, et al. Stents and surgical interventions in the palliation of gastric outlet obstruction: a systematic review. Endosc Int Open 2016;4:E1158-70.
- Mintziras I, Miligkos M, Wachter S, et al. Palliative surgical bypass is superior to palliative endoscopic stenting in patients with malignant gastric outlet obstruction: systematic review and meta-analysis. Surg Endosc 2019;33:3153-64.
- Nagaraja V, Eslick GD, Cox MR. Endoscopic stenting versus operative gastrojejunostomy for malignant gastric outlet obstruction—a systematic review and meta-analysis of randomized and nonrandomized trials. J Gastrointest Oncol 2014;5:92-8.
- Upchurch E, Ragusa M, Cirocchi R. Stent placement versus surgical palliation for adults with malignant gastric outlet obstruction. Cochrane Database System Rev 2018;30:CD012506.
- 24. Zheng B, Wang X, Ma B, et al. Endoscopic stenting versus gastrojejunostomy for palliation of malignant gastric outlet obstruction. Dig Endosc 2012;24:71-8.
- 25. Fiori E, Lamazza A, Demasi E, et al. Endoscopic stenting for gastric outlet obstruction in patients with unresectable antro pyloric cancer. Systematic review of the literature and final results of a prospective study. The point of view of a surgical group. Am J Surg 2013;206:210-7.
- Jeurnink S, Steyerberg E, van Hooft JE, et al. Surgical gastrojejunostomy or endoscopic stent placement for the palliation of malignant gastric outlet obstruction (SUSTENT study): a multicenter randomized trial. Gastrointest Endosc 2010;71:490-9.
- Mehta S, Hindmarsh A, Cheong E, et al. Prospective randomized trial of laparoscopic gastrojejunostomy versus duodenal stenting for malignant gastric outflow obstruction. Surg Endosc 2006;20:239-42.
- Sterne JA, Hernan MA, Reeves BC, et al. ROBINS-I: a tool for assessing risk of bias in non-randomised studies of interventions. BMJ 2016;355L:i4919.
- Dindo D, Demartines N, Clavien PA. Classification of surgical complications: a new proposal with evaluation in a cohort of 6336 patients and results of a survey. Ann Surg 2004;240:205-13.
- Chandrasegaram MD, Eslick GD, Mansfield CO, et al. Endoscopic stenting versus operative gastrojejunostomy for malignant gastric outlet obstruction. Surg Endosc 2012;26:323-9.
- El-Shabrawi A, Cerwenka H, Bacher H, et al. Treatment of malignant gastric outlet obstruction: endoscopic implantation of self-expanding metal stents versus gastric bypass surgery. Eur Surg 2006;38:451-5.
- 32. Leiyuan S, Jianli X, Zhengzhong Z, et al. Comparison of treatment outcomes of endoscopic stenting and laparoscopic gastrojejunostomy for malignant gastric outlet obstruction. Am Surg 2018;84:991-5.
- Min SH, Son SY, Jung DH, et al. Laparoscopic gastrojejunostomy versus duodenal stenting in unresectable gastric cancer with gastric outlet obstruction. Ann Surg Treat Res 2017;93:130-6.
- 34. No JH, Kim SW, Lim CH, et al. Long-term outcome of palliative therapy for gastric outlet obstruction caused by unresectable gastric cancer in patients with good performance status: endoscopic stenting versus surgery. Gastrointest Endosc 2013;78:55-62.
- Rudolph HU, Post S, Schluter M, et al. Malignant gastroduodenal obstruction: retrospective comparison of endoscopic and surgical palliative therapy. Scand J Gastroenterol 2011;46:583-90.
- Yoshida Y, Fukutomi A, Tanaka M, et al. Gastrojejunostomy versus duodenal stent placement for gastric outlet obstruction in patients with unresectable pancreatic cancer. Pancreatology 2017;17:983-9.
- Ware JE Jr, Gandek B. Overview of the SF-36 Health Survey and the International Quality of Life Assessment (IQOLA) project. J Clin Epidemiol 1998;51:903-12.
- **38.** Jang SH, Lee H, Min BH, et al. Palliative gastrojejunostomy versus endoscopic stent placement for gastric outlet obstruction in patients with unresectable gastric cancer: a propensity score-matched analysis. Surg Endosc 2017;31:4217-23.

- **39.** Jeurnink SM, Steyerberg EW, Vleggaar FP, et al. Predictors of survival in patients with malignant gastric outlet obstruction: a patient-oriented decision approach for palliative treatment. Dig Liver Dis 2011;43:548-52.
- **40.** Siddiqui A, Spechler SJ, Huerta S. Surgical bypass versus endoscopic stenting for malignant gastroduodenal obstruction: a decision analysis. Dig Dis Sci 2007;52:276-81.
- Fisher AV, Hanlon B, Fernandes-Taylor S, et al. Natural history and cost analysis of surgical bypass versus endoscopic stenting for the palliative management of malignant gastric outlet obstruction. HPB (Oxford) 2020;22:529-36.
- **42.** Jeurnink SM, Polinder S, Steyerberg EW, et al. Cost comparison of gastrojejunostomy versus duodenal stent placement for malignant gastric outlet obstruction. J Gastroenterol 2010;45:537-43.
- **43.** Kim CG, Park SR, Choi IJ, et al. Effect of chemotherapy on the outcome of self-expandable metallic stents in gastric cancer patients with malignant outlet obstruction. Endoscopy 2012;44:807-12.
- 44. Kobayashi S, Ueno M, Kameda R, et al. Duodenal stenting followed by systemic chemotherapy for patients with pancreatic cancer and gastric outlet obstruction. Pancreatology 2016;16:1085-91.
- **45.** Jang S, Stevens T, Lopez R, et al. Superiority of gastrojejunostomy over endoscopic stenting for palliation of malignant gastric outlet obstruction. Clin Gastroenterol Hepatol 2019;17:1295-302.
- **46.** Chen YI, Kunda R, Storm AC, et al. EUS-guided gastroenterostomy: a multicenter study comparing the direct and balloon-assisted techniques. Gastrointest Endosc 2018;87:1215-21.
- 47. McCarty TR, Garg R, Thompson CC, et al. Efficacy and safety of EUS-guided gastroenterostomy for benign and malignant gastric outlet obstruction: a systematic review and meta-analysis. Endosc Int Open 2019;7:E1474-82.
- **48.** Ge PS, Young JY, Dong W, et al. EUS-guided gastroenterostomy versus enteral stent placement for palliation of malignant gastric outlet obstruction. Surg Endosc 2019;33:3404-11.
- **49.** Chen YI, Itoi T, Baron TH, et al. EUS-guided gastroenterostomy is comparable to enteral stenting with fewer re-interventions in malignant gastric outlet obstruction. Surg Endosc 2017;31:2946-52.
- 50. Khashab MA, Bukhari M, Baron TH, et al. International multicenter comparative trial of endoscopic ultrasonography-guided gastroenterostomy versus surgical gastrojejunostomy for the treatment of malignant gastric outlet obstruction. Endosc Int Open 2017;5: E275-81.
- Hamada T, Hakuta R, Takahara N, et al. Covered versus uncovered metal stents for malignant gastric outlet obstruction: systematic review and meta-analysis. Dig Endosc 2017;29:259-71.
- Pan YM, Pan J, Guo LK, et al. Covered versus uncovered selfexpandable metallic stents for palliation of malignant gastric outlet obstruction: a systematic review and meta-analysis. BMC Gastroenterol 2014;14:170.
- 53. Van Halsema EE, Rauws EAJ, Fockens P, et al. Self-expandable metal stents for malignant gastric outlet obstruction: a pooled analysis of prospective literature. World J Gastroenterol 2015;21:12468-81.
- 54. Kim C, Choi I, Lee J, et al. Covered versus uncovered self-expandable metallic stents for palliation of malignant pyloric obstruction in gastric cancer patients: a randomized, prospective study. Gastrointest Endosc 2010;72:25-32.
- 55. Maetani I, Mizumoto Y, Shigoka H, et al. Placement of a triple-layered covered versus uncovered metallic stent for palliation of malignant gastric outlet obstruction: a multicenter randomized trial. Dig Endosc 2014;26:192-9.
- 56. Lee H, Min BH, Lee JH, et al. Covered metallic stents with an antimigration design vs. uncovered stents for the palliation of malignant gastric outlet obstruction: a multicenter, randomized trial. Am J Gastroenterol 2015;110:1440-9.
- 57. Lim S, Kim J, Lee K, et al. Conformable covered versus uncovered self-expandable metallic stents for palliation of malignant gastroduodenal obstruction: a randomized prospective study. Dig Liver Dis 2014;46:603-8.

- 58. Shi D, Ji F, Bao YS, et al. A multicenter randomized controlled trial of malignant gastric outlet obstruction: tailored partially covered stents (placed fluoroscopically) versus standard uncovered stents (placed endoscopically). Gastroenterol Res Pract 2014:309797.
- Bang S, Kim HJ, Park JY, et al. Effectiveness of self-expanding metal stents for malignant antropyloric and duodenal obstruction with a comparison between covered and uncovered stents. Hepatogastroenterology 2008;55:2091-5.
- 60. Hori Y, Naitoh I, Hayashi K, et al. Predictors of stent dysfunction after self-expandable metal stent placement for malignant gastric outlet obstruction: tumor ingrowth in uncovered stents and migration of covered stents. Surg Endosc 2017;31:4165-73.
- 61. Jung K, Ahn JY, Jung HY, et al. Outcomes of endoscopically inserted self-expandable metal stents in malignancy according to the type of stent and the site of obstruction. Surg Endosc 2016;30:4001-10.
- Kim JW, Jeong JB, Lee KL, et al. Comparison between uncovered and covered self-expandable metal stent placement in malignant duodenal obstruction. World J Gastroenterol 2015;21: 1580-7.
- 63. Lee KM, Choi SJ, Shin SJ, et al. Palliative treatment of malignant gastroduodenal obstruction with metallic stent: prospective comparison of covered and uncovered stents. Scand J Gastroenterol 2009;44: 846-52.
- Maetani I, Ukita T, Tada T, et al. Metallic stents for gastric outlet obstruction: reintervention rate is lower with uncovered versus covered stents, despite similar outcomes. Gastrointest Endosc 2009;69:806-12.
- Park CI, Kim JH, Lee YC, et al. What is the ideal stent as initial intervention for malignant gastric outlet obstruction? Dig Liver Dis 2013;45:33-7.
- 66. Woo SM, Kim DH, Lee WJ, et al. Comparison of uncovered and covered stents for the treatment of malignant duodenal obstruction caused by pancreaticobiliary cancer. Surg Endosc 2013;27:2031-9.
- Maetani I. Self-expandable metallic stent placement for palliation in gastric outlet obstruction. Ann Palliat Med 2014;3:54-64.
- Ge PS, Watson RR, Chen DC, et al. Delayed migration of a WallFlex enteral stent resulting in jejunal perforation. Case Rep Gastrointest Med 2013;2013:65259.
- Tringali A, Giannetti A, Adler DG. Endoscopic management of gastric outlet obstruction disease. Ann Gastroenterol 2019;32:330-7.
- Hori Y, Hayashi K, Naitoh I, et al. Feasibility and safety of duodenal covered self-expandable metallic stent fixation: an experimental study. Surg Endosc 2019;33:4026-31.
- Kim ID, Kang DH, Choi CW, et al. Prevention of covered enteral stent migration in patients with malignant gastric outlet obstruction: a pilot study of anchoring with endoscopic clips. Scand J Gastroenterol 2010;45:100-5.
- Abdel-Salam WN, Katri KM, Bessa SS, et al. Laparoscopic-assisted truncal vagotomy and gastro-jejunostomy: trial of simplification. J Laparoendosc Adv Surg Tech A 2009;19:125-7.
- Jani K, Saxena A, Kothari A. Laparoscopic truncal vagotomy with gastrojejunostomy. J Indian Med Assoc 2010;108:648-51.
- 74. Kim SM, Song J, Oh SJ, et al. Comparison of laparoscopic truncal vagotomy with gastrojejunostomy and open surgery in peptic pyloric stenosis. Surg Endosc 2009;23:1326-30.
- Palanivelu C, Jani K, Rajan PS, et al. Laparoscopic management of acid peptic disease. Surg Laparosc Endosc Percutan Tech 2006;16: 312-6.
- Rangarajan M, Subramanian CS, Chandralathan TA. Laparoscopyassisted truncal vagotomy with antecolic posterior gastrojejunostomy for benign gastric outlet obstruction. Surg Endosc 2006;20: 61-3.
- Siu WT, Tang CN, Law BK, et al. Vagotomy and gastrojejunostomy for benign gastric outlet obstruction. J Laparoendosc Adv Surg Tech A 2004;14:266-9.
- Soreide K, Sarr MG, Soreide JA. Pyloroplasty for benign gastric outlet obstruction—indications and techniques. Scand J Surg 2006;95:11-6.

- Boylan JJ, Gradzka MI. Long-term results of endoscopic balloon dilatation for gastric outlet obstruction. Dig Dis Sci 1999;44:1883-6.
- **80.** Cherian PT, Cherian S, Singh P. Long-term follow-up of patients with gastric outlet obstruction related to peptic ulcer disease treated with endoscopic balloon dilatation and drug therapy. Gastrointest Endosc 2007:66:491-7.
- **81.** Chiu YC, Liang CM, Tam W, et al. The effects of endoscopic-guided balloon dilations in esophageal and gastric strictures caused by corrosive injuries. BMC Gastroenterol 2013;13:99.
- **82.** DiSario JA, Fennerty MB, Tietze CC, et al. Endoscopic balloon dilation for ulcer-induced gastric outlet obstruction. Am J Gastroenterol 1994;89:868-71.
- Hamzaoui L, Bouassida M, Ben Mansour I, et al. Balloon dilatation in patients with gastric outlet obstruction related to peptic ulcer disease. Arab J Gastroenterol 2015;16:121-4.
- 84. Hewitt PM, Krige JE, Funnell IC, et al. Endoscopic balloon dilatation of peptic pyloroduodenal strictures. J Clin Gastroenterol 1999;28:33-5.
- Kochhar R, Dutta U, Sethy PK, et al. Endoscopic balloon dilation in caustic-induced chronic gastric outlet obstruction. Gastrointest Endosc 2009:69:800-5.
- **86.** Kochhar R, Malik S, Gupta P, et al. Etiological spectrum and response to endoscopic balloon dilation in patients with benign gastric outlet obstruction. Gastrointest Endosc 2018;88:899-908.
- 87. Kochhar R, Sethy PK, Nagi B, et al. Endoscopic balloon dilatation of benign gastric outlet obstruction. J Gastroenterol Hepatol 2004;19:418-22.
- 88. Lam YH, Yun-wong Lau J, Ming-kit Fung T, et al. Endoscopic balloon dilation for benign gastric outlet obstruction with or without *Helicobacter pylori* infection. Gastrointest Endosc 2004;60:229-33.
- **89.** Lau JY, Chung SC, Sung JJ, et al. Through-the-scope balloon dilation for pyloric stenosis: long-term results. Gastrointest Endosc 1996;43(2 Pt 1):98-101.
- Noor MT, Dixit P, Kochhar R, et al. NSAIDs-related pyloroduodenal obstruction and its endoscopic management. Diagn Ther Endosc 2011;2011:967957.
- 91. Rana SS, Bhasin DK, Chandail VS, et al. Endoscopic balloon dilatation without fluoroscopy for treating gastric outlet obstruction because of benign etiologies. Surg Endosc 2011;25:1579-84.
- 92. Choi WJ, Park JJ, Park J, et al. Effects of the temporary placement of a self-expandable metallic stent in benign pyloric stenosis. Gut Liver 2013;7:417-22.
- 93. Heo J, Jung MK. Safety and efficacy of a partially covered selfexpandable metal stent in benign pyloric obstruction. World J Gastroenterol 2014;20:16721-5.
- 94. Bazerbachi F, Heffley JD, Abu Dayyeh BK, et al. Safety and efficacy of coaxial lumen-apposing metal stents in the management of refractory gastrointestinal luminal strictures: a multicenter study. Endosc Int Open 2017;5:E861-7.
- **95.** Jain D, Patel U, Ali S, et al. Efficacy and safety of lumen-apposing metal stent for benign gastrointestinal stricture. Ann Gastroenterol 2018:31:425-38.
- Larson B, Adler DG. Lumen-apposing metal stents for gastrointestinal luminal strictures: current use and future directions. Ann Gastroenterol 2019;32:141-6.
- Santos-Fernandez J, Paiji C, Shakhatreh M, et al. Lumen-apposing metal stents for benign gastrointestinal tract strictures: an international multicenter experience. World J Gastrointest Endosc 2017;9:571-8.
- **98.** Yang D, Nieto JM, Siddiqui A, et al. Lumen-apposing covered self-expandable metal stents for short benign gastrointestinal strictures: a multicenter study. Endoscopy 2017;49:327-33.
- **99.** Khan K. High-resolution EUS to differentiate hypertrophic pyloric stenosis. Gastrointest Endosc 2008;67:375-6.

- 100. Sharma D, Bharany RP, Mapshekhar RV. Duplication cyst of pyloric canal: a rare cause of pediatric gastric outlet obstruction: rare case report. Indian J Surg 2013;75(Suppl 1):322-5.
- 101. Chao HC. Update on endoscopic management of gastric outlet obstruction in children. World J Gastrointest Endosc 2016;8: 635-45.
- 102. Dehghani SM, Aldaghi M, Javaherizadeh H. Endoscopic pyloroplasty for severe gastric outlet obstruction due to alkali ingestion in a child. Gastroenterol Hepatol Bed Bench 2016;9:64-7.
- 103. Fishman DS, Kellermayer R, Lopez M, et al. Pediatric usage of a biliary endoprosthesis for gi and biliary disease [abstract]. Gastrointestinal Endoscopy 2013;77:AB244-5.
- 104. Ibarguen-Secchia E. Endoscopic pyloromyotomy for congenital pyloric stenosis. Gastrointest Endosc 2005;61:598-600.
- 105. Ricciuto A, Connolly BL, Gonska T. Serial balloon dilation to relieve gastric outlet obstruction induced by the ingestion of toilet cleaner. J Pediatr Gastroenterol Nutr 2018;66:e56.

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UPPLEMEN	ITARY TABLE	1. Systen	natic revi	ew of cov	ered versus	uncovered S	EMS adverse	events			
Reference	Type of study	Covered	Un- covered	Bleeding covered	Bleeding uncovered		Perforation uncovered	Cholangitis covered	Cholangitis uncovered	Pancreatitis covered	Pancreatiti uncovered
Shi et al, 2014 ⁵⁸	RCT	33	32	11*	2	0	0	N	N	N	N
Lim et al, 2014 ⁵⁷	RCT	66	68	0	0	0	0	N	N	N	N
Kim et al, 2010 ⁵⁴	RCT	40	40	0	0	1	0	N	N	N	N
Maetani et al, 2014 ⁵⁵	RCT	31	31	0	1	1	0	N	N	N	N
Lee et al, 2015 ⁵⁶	RCT	51	51	N	N	N	N	N	N	N	N
Hori et al, 2017 ⁶⁰	Retrospective	126	126	1	0	3	0	1	1	1	1
Jung et al, 2016 ⁶¹	Retrospective	93†	120	N	N	N	N	N	N	N	N
Kim et al, 2015 ⁶²	Retrospective	29	38	N	N	3	0	N	N	N	N
Park et al, 2013 ⁶⁵	Retrospective	64	128	N	N	N	N	N	N	N	N
Woo et al, 2013 ⁶⁶	Retrospective	24	46	1	1	2	2	N	N	N	N
Maetani et al, 2009 ⁶⁴	Retrospective	29	31	0	1	0	1	N	N	1	1
Bang et al, 2008 ⁵⁹	Retrospective	53	79	N	N	N	N	N	N	N	N
Lee et al, 2009 ⁶³	Prospective	70	84	0	0	0	0	N	N	N	N
Totals		709	874	2	5	10	3	1	1	2	2
				2/386	5/458	10/448	3/496	1/126	1/126	2/155	2/157
Event rate,				0.5	1.1	2.2	0.6	0.8	0.8	1.3	1.3

SEMS, Self-expandable metal stent; RCT, randomized controlled trial; N, not available.

 $[\]hbox{*Funnel-shaped covered SEMS and outlier omitted from bleeding calculation.}$

[†]Seventy-seven partially covered, 16 fully covered.

SUPPLEMENTARY TABLE 2. Systematic review of successful endoscopic balloon dilation treatment of benign gastric outlet obstruction caused by NSAIDs, *Helicobacter pylori*, and pancreatitis

Reference	No. of patients	NSAIDs	Helicobacter pylori	Pancreatitis
Boylan & Gradzka, 1999 ⁷⁹	40	19/24 (79.1)		
Cherian et al, 2007 ⁸⁰	23	8/8 (100)	17/17 (100)	
Chiu et al, 2013 ⁸¹	7			
DiSario et al, 1994 ⁸²	30	12/15 (80)		
Misra et al, 1998 ³	14			
Hamzaoui et al, 2015 ⁸³	45			
Hewitt et al, 1999 ⁸⁴	41			
Kochhar et al, 2009 ⁸⁵	41*			
Kochhar et al, 2018 ⁸⁶	264	97.8†		
Kochhar et al, 2004 ⁸⁷	23			0/4 (0)
Lam et al, 2004 ⁸⁸	33		11/14 (79)	
Lau et al, 1996 ⁸⁹	41			
Noor et al, 2011 ⁹⁰	10	9/10 (90)		
Rana et al, 2011 ⁹¹	25		10/11 (91)‡	3/5 (60)

Values are n/N (%).

NSAID, Nonsteroidal anti-inflammatory drug.

^{*}Data overlaps with 2018 study by same author.

 $[\]dagger$ Patients on opioids (n = 10) and on opioids plus NSAIDs (n = 12) were grouped together.

[‡]Helicobacter pylori patients were grouped together with 2 patients who took NSAIDs.

APPENDIX 1

Search strategy

Search date: September 6, 2018

Databases searched: Ovid MEDLINE: Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE Daily, and Ovid MEDLINE 1946 to Present, Embase Classic+Embase 1947 to 2018 September 6; Wiley Cochrane

Population, Intervention, Comparator, and Outcomes 1 and 2

Ovid MEDLINE(R), Embase

Number	Searches	Results
1	exp Gastric Outlet Obstruction/ use ppez	5550
2	exp stomach obstruction/ use emczd	3981
3	((gastric or GI or gastrointestinal or gastroduodenal) adj4 obstruction).ti,ab.	9462
4	goo.ti,ab.	734
5	or/1-4	15,535
6	exp Stents/ use ppez	69,495
7	exp stent/ use emczd	153,959
8	(stent* or sems).ti,ab.	237,684
9	(EUS-GJ or (endoscop* adj3 gastrojejunostomy)).ti,ab.	330
10	(EUS-GE or (endoscop* adj3 gastroenterostomy)).ti,ab.	72
11	or/6-10	286,140
12	5 and 11	1921
13	limit 12 to english language	1738
14	animals/ not (humans/ and animals/)	5,782,738
15	13 not 14	1729
16	remove duplicates from 15	1231
17	Meta - Analysis/ use ppez or Meta - Analysis as Topic/ use ppez or exp Technology Assessment, Biomedical/ use ppez	117,957
18	Meta Analysis/ use emczd or "Meta Analysis (Topic)"/ use emczd or Biomedical Technology Assessment/ use emczd	194,284
19	Meta Analysis.pt.	92,260
20	(meta analy* or metaanaly* or health technolog* assess*).mp.	410,060
21	(((systematic* or methodologic*) adj3 (review* or overview*)) or pooled analysis or published studies or published literature or hand search* or handsearch* or medline or pub med or pubmed or embase or cochrane or cinahl or data synthes* or data extraction* or HTA or HTAs or (technolog* adj (assessment* or overview* or appraisal*))).ti,ab.	641,188

. Continued		
Number	Searches	Results
22	or/17-21	847,523
23	16 and 22	51
24	remove duplicates from 23	51

Wiley Cochrane

ID	Search	hits
#1	MeSH descriptor: [Gastric Outlet Obstruction] explode all trees	65
#2	((gastric or GI or gastrointestinal or gastroduodenal) near/4 obstruction):ti,ab	175
#3	goo:ti,ab	39
#4	#1 or #2 or #3	226
#5	MeSH descriptor: [Stents] explode all trees	3806
#6	(stent* or SEMS):ti,ab	10,644
#7	(EUS-GJ or (endoscop* NEAR/3 gastrojejunostomy)):ti,ab	13
#8	(EUS-GE or (endoscop* NEAR/3 gastroenterostomy)):ti,ab	4
#9	#5 or #6 or #7 or #8	10,951
#10	#4 and #9	49

Population, Intervention, Comparator, and Outcome 3

Ovid MEDLINE(R), Embase

Number	Searches	Results
1	exp Gastric Outlet Obstruction/ use ppez	5550
2	exp stomach obstruction/ use emczd	3981
3	((gastric or GI or gastrointestinal or gastroduodenal) adj4 obstruction).ti,ab.	9462
4	goo.ti,ab.	734
5	or/1-4	15,535
6	exp Dilatation/ use ppez	10,560
7	exp balloon dilatation/ use emczd	17,270
8	(dilatation or dilation).ti,ab.	213,935
9	or/6-8	227,722
10	5 and 9	1010
11	limit 10 to english language	910
12	animals/ not (humans/ and animals/)	5,782,738
13	11 not 12	891
	(continued on	the next page)

. Continu	. Continued			
Number	Searches	Results		
14	remove duplicates from 13	703		
15	Meta - Analysis/ use ppez or Meta - Analysis as Topic/ use ppez or exp Technology Assessment, Biomedical/ use ppez	117,957		
16	Meta Analysis/ use emczd or "Meta Analysis (Topic)"/ use emczd or Biomedical Technology Assessment/ use emczd	194,284		
17	Meta Analysis.pt.	92,260		
18	(meta analy* or metaanaly* or health technolog* assess*).mp.	410,060		
19	(((systematic* or methodologic*) adj3 (review* or overview*)) or pooled analysis or published studies or published literature or hand search* or handsearch* or medline or pub med or pubmed or embase or cochrane or cinahl or data synthes* or data extraction* or HTA or HTAs or (technolog* adj (assessment* or overview* or appraisal*))).ti,ab.	641,188		
20	or/15-19	847,523		
21	14 and 20	12		
22	remove duplicates from 21	12		

Wiley Cochrane

ID	Search	hits
#1	MeSH descriptor: [Gastric Outlet Obstruction] explode all trees	65
#2	((gastric or GI or gastrointestinal or gastroduodenal) near/4 obstruction):ti,ab	175
#3	goo:ti,ab	39
#4	#1 or #2 or #3	226
#5	MeSH descriptor: [Dilatation] explode all trees	389
#6	(dilatation or dilation):ti,ab	6818
#7	#5 or #6	6958
#8	#4 and #7	5