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META-ANALYSIS

Endoscopic ultrasound-guided vs endoscopic retrograde cholangiopancreatography biliary drainage for obstructed distal malignant biliary strictures: A systematic review and meta-analysis

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Abstract

BACKGROUND

For palliation of malignant biliary obstruction (MBO), the gold-standard method of biliary drainage is endoscopic retrograde cholangiopancreatography (ERCP) with the placement of metallic stents. Endoscopic ultrasound (EUS)-guided drainage is an alternative that is typically reserved for cases of ERCP failure. Recently, however, there have been robust randomized clinical trials (RCTs) comparing EUS-guided drainage and ERCP as primary approaches to MBO.

AIM

To compare EUS guidance and ERCP in terms of their effectiveness and safety in palliative biliary drainage for MBO.

METHODS

This was a systematic review and meta-analysis, in which we searched the MEDLINE, *Excerpta Medica*, and Cochrane Central Register of Controlled Trials databases. Only RCTs comparing EUS and ERCP for primary drainage of MBO were eligible. All of the studies selected provided data regarding the rates of technical and clinical success, as well as the duration of the procedure, adverse

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events, and stent patency. We assessed the risk of biases using the Jadad score and the quality of evidence using the Grading of Recommendations Assessment, Development and Evaluation criteria.

RESULTS

The database searches yielded 5920 records, from which we selected 3 RCTs involving a total of 222 patients (112 submitted to EUS and 110 submitted to ERCP). In the EUS and ERCP groups, the rate of technical success was 91.96% and 91.81%, respectively, with a risk difference (RD) of 0.00% (95%CI: -0.07, 0.07; P = 0.97; P = 0

CONCLUSION

EUS represents an interesting alternative to ERCP for MBO drainage, demonstrating lower stent dysfunction rates compared with ERCP. Technical and clinical success, duration, adverse events and patency rates were similar.

Key words: : Common bile duct neoplasms; Endoscopic retrograde cholangiopancreatography; Endosonography; Ultrasonography; Interventional/methods; Endoscopic ultrasound; Systematic review; Meta-analysis

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Core tip: No consensus is available in the literature regarding whether endoscopic retrograde cholangiopancreatography or endoscopic ultrasound-guided biliary drainage is more beneficial to the patient. This is the first systematic review and meta-analysis comparing the two methods. We investigated these two techniques in terms of technical and clinical success, as well as duration of the procedure, adverse events, stent dysfunction and stent patency.

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INTRODUCTION

Endoscopic retrograde cholangiopancreatography (ERCP) is currently the goldstandard method to address malignant biliary obstruction (MBO) of the distal common bile duct^[1,2], the procedure consists in endoscopic guidewire access to the duodenal papilla, with further injection of contrast on the bile ducts and placement of an endoscopic stent in order to treat MBO, and there are data favoring the use of selfexpanding metal stents over that of plastic stents^[3]. However, ERCP is not free of complications, the most common being post-ERCP pancreatitis and cholangitis^[4]. In addition, there is a non-negligible risk of failed biliary cannulation in ERCP due to dysfunctional biliary sphincter or anatomical alterations^[5].

Percutaneous transhepatic biliary drainage (PTBD) and surgical bilioenteric anastomosis are traditional alternatives to ERCP, although both have their particular drawbacks. PTBD requires multiple interventions and carries an increased risk of cholangitis, bacteremia, and hemobilia^[6], whereas bilioenteric anastomosis is associated with high morbidity and mortality^[7].

Endoscopic ultrasound (EUS) has long been of paramount importance for the



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workup of patients with biliary obstruction^[8-10]. Some recent reports have described EUS-guided drainage as an alternative in cases of ERCP failure^[11-13]. The efficacy and safety profile of EUS-guided drainage have improved over time, as has the availability of specific accessories, allowing some authors to test EUS-guided biliary drainage, in comparison with ERCP, as a primary approach to biliary obstruction.

Transluminal EUS-guided biliary drainage consists of needle access to the biliary ducts by hepatogastric or choledocoduodenal puncture under EUS guidance. Then, a guidewire is inserted through the needle, followed by dilation of the fistula and stent placement.

Although there have been a number of randomized clinical trials (RCTs) comparing EUS-guided biliary drainage and ERCP^[14-16], there have yet to be any systematic reviews or meta-analyses regarding the topic. Therefore, the aim of the present study was to summarize all available data comparing EUS and ERCP in terms of their effectiveness and safety in the primary drainage of MBO. To that end, we conducted a systematic review and meta-analysis, in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) methodology, of RCTs comparing EUS and ERCP in the primary drainage of distal MBO, assessing technical success, clinical success, cost-effectiveness, duration of the procedure, adverse events, mortality, stent patency, and stent dysfunction.

MATERIALS AND METHODS

Protocol and registration

This study followed the PRISMA guidelines^[17] and was registered in the International Prospective Register of Systematic Reviews database^[18] (CRD42018108712). The study was approved by the local institutional review board.

Study criteria

Only RCTs were considered eligible, without barriers as to the language or year of publication. We included RCTs that had evaluated patients diagnosed with distal MBO and undergoing primary drainage of the biliary tract under EUS guidance or by ERCP. Studies evaluating patients with benign biliary obstruction were excluded, as were those evaluating EUS-guided biliary drainage after failure of another method and those including only patients undergoing primary EUS-guided drainage due to an anatomical alteration that precluded ERCP.

Search strategy and study selection

We searched the MEDLINE, Excerpta Medica, Cochrane Central Register of Controlled Trials, Latin-American and Caribbean Health Sciences Literature databases, as well as the gray literature, for RCTs published up to and including November 2018. We employed descriptors available from the United States National Library of Medicine Medical Subject Headings and, to a lesser degree, other related terms aiming at a more sensitive strategy. For Medline, our search strategy was as follows: [(ERCP OR Endoscopic Retrograde Cholangiopancreatograph*) OR (EUS OR endosonography OR Endoscopic Ultrasonograph* OR Echo Endoscop*)] AND (decompression OR drain*). For the other databases, the following search strategy was applied: (EUS OR Endoscopic Ultrasonography) AND (decompression OR drainage).

Two independent researchers assessed titles and abstracts for eligibility. Any disagreement was resolved by consensus with a third experienced researcher. The articles were included after an evaluation of the full-text based on the study criteria.

Data extraction and evaluation

Data related to EUS-guided and ERCP biliary drainage were collected using a preformatted Excel workbook. The data collected included technical and clinical success rates, as well as the duration of the procedure, adverse events, stent patency, and stent dysfunction.

In our quantitative analysis, we used the absolute values, means, and standard deviations. If a study expressed outcomes using median and interquartile range, mathematical formulas were used for data conversion^[19]. In case of the study fails to present means and standard deviations or median and interquartile range of the continuous variables of specific outcomes, rendering impossible to include the data for meta-analysis evaluation, the study in question was excluded from the outcome appraisal.

Evaluation of biases and quality of studies

The biases of the RCTs were assessed with the Jadad scale^[20], which allows critical appraisal regarding blinding, randomization, and information on losses to follow-up.



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Jadad score is applied to evaluate the methodological quality of RCTs, rating the study from zero (poor quality) to five points (rigorous). The evaluation criteria are "description of the study as randomized", "employment of appropriated randomization method", "description of the method of blinding", "employment of appropriated blinding method" and "description of losses to follow-up" whereupon each present criteria grants one point.

The quality of evidence was assessed using the Grading of Recommendations Assessment, Development and Evaluation (GRADE) criteria with the GRADEpro Guideline Development Tool software (McMaster University, 2015; Evidence Prime, Inc., Ontario, Canada)^[21]. GRADE is an approach to rate the quality of evidence based on criteria guideline developed by the GRADE working group and involves the appraisal of risk of bias, inconsistency, indirectness, imprecision and publication bias. Evaluation of biases and quality of studies was performed under supervision of our statistic team.

Data analysis

For the dichotomous variables, we calculated the risk difference (RD) values, using the Mantel-Haenszel test, together with the corresponding 95% confidence intervals. For continuous variables, we calculated the mean difference values, also with the corresponding 95% confidence intervals, using the inverse variance test. The results were displayed with forest plots.

We assessed the heterogeneity among studies using the Higgins test (l^2). If there was an $l^2 < 50\%$, we used a fixed-effect model, whereas we used funnel plot analysis if there was an $l^2 > 50\%$. If we detected an outlier article, we removed it from the analysis and kept the fixed-effect model. If we could not detect an outlier, we switched to the random-effect model analysis to ameliorate the impact of the high heterogeneity. All analyses were carried out with Review Manager software, version 5.3.5 (RevMan 5; Cochrane Collaboration, Oxford, United Kingdom).

RESULTS

Overview

The database searches retrieved a total of 5920 studies, 164 of which were selected for full-text evaluation. Based on the study criteria, three RCTs were included in the qualitative analysis and meta-analysis (Figure 1).

The collective sample comprised 222 patients: 112 in the EUS group and 110 in the ERCP group. The mean age was similar between the two groups and among the samples of the RCTs included. The etiology of MBO in the studies selected is outlined in Table 1.

On the Jadad scale (Table 2), all of the RCTs evaluated had a score of 3, which is the highest possible score for unblinded studies. According to the GRADE criteria for the quality of evidence, the evidence for technical success generated moderate certainty, the evidence for stent dysfunction generated low certainty, and the evidence for the remaining outcomes generated very low certainty (Table 3).

Technical success

All three RCTs^[14-16] reported technical success rates. The mean rate of technical success was 91.96% and 91.81% in the EUS and ERCP groups, respectively, with an RD of 0.00% (95%CI: -0.07, 0.07; P = 0.97), demonstrating no statistical difference between the two techniques (Figure 2).

Clinical success

All three RCTs included data on clinical success^[14-16]. However, Bang *et al*^[16] included cross-over procedures in their final results, precluding the intention-to-treat analysis and thus excluding 67 patients. Therefore, the final collective sample in our analysis of clinical success comprised 155 patients: 79 in the EUS group and 76 in the ERCP group. The mean clinical success rate was 84.81% and 85.53% in the EUS and ERCP groups, respectively, with an RD of -0.01% (95%CI: -0.12, 0.10; P = 0.90), as shown in Figure 3.

Duration of the procedure

All three studies^[14-16] reported the duration of the procedure in minutes. The mean time difference between EUS-guided and ERCP drainage was -0.12% (95%CI: -8.20, 7.97; P = 0.98), showing no statistical difference between the two groups (Figure 4). We found high heterogeneity among the studies ($I^2 = 84\%$). Because there were no outliers, we employed the random-effect model in our analysis.

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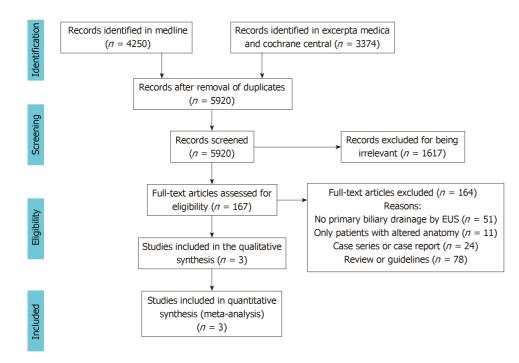


Figure 1 Flow chart of study selection. Cochrane CENTRAL: Cochrane Central Register of Controlled Trials; EUS: Endoscopic ultrasound.

Adverse events

All three RCTs^[14-16] described the adverse events reported. In the EUS group, 14 adverse events were reported: abdominal pain (n = 5); cholangitis (n = 4); pneumoperitoneum (n = 2); biliary peritonitis (n = 2); and cholecystitis (n = 1). In the ERCP group, there were 25 adverse events: pancreatitis (n = 10); cholangitis (n = 7); cholecystitis (n = 5); and abdominal pain (n = 3). No procedure-related mortality was reported in any of the studies.

Although we identified high heterogeneity ($l^2 = 77\%$), there were no outlier studies, and the random-effect model was therefore employed. The mean difference between the two techniques was -0.06% (95%CI: -0.23, 0.12; P = 0.54), indicating that there was no statistical difference (Figure 5).

Stent patency

Although all three RCTs reported data on stent patency, Paik *et al*^[15] did not detail standard deviation values, precluding the inclusion of that study in the analysis and thus excluding 125 patients. Therefore, the final collective sample in our analysis of stent patency comprised 97 patients: 48 in the EUS group and 49 in the ERCP group. The mean difference was 9.32% (95%CI: -4.53, 23.18; P = 0.19), demonstrating no significant difference between the two methods in terms of stent patency (Figure 6)

Stent dysfunction

All three studies^[14-16] provided data on stent dysfunction. We found high heterogeneity ($l^2 = 86\%$) among the studies, and the funnel plot analysis identified the Bang *et al*^[16] study as an outlier. When we excluded that study from the analysis, the l^2 value was 0%. Between the two remaining studies, there were 12 cases of stent dysfunction requiring intervention in the EUS group and 28 such cases in the ERCP group. The RD between the groups was -0.22% (95%CI: -0.35, -0.08; P = 0.001), thus favoring EUS-guidance over ERCP (Figure 7).

DISCUSSION

Summary of evidence

To our knowledge, this is the first systematic review and meta-analysis including only RCTs that compared EUS-guidance and ERCP as the primary approach to biliary drainage in cases of MBO. Our strict methodology, which included critical appraisal of biases, quality of evidence assessment, and a report prepared in accordance with the PRISMA guidelines^[17], underscores the strength of our findings.

EUS-guided biliary drainage was first introduced as an alternative to be employed



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Table 1 Characteristics of the patient samples and etiology of malignant biliary obstruction in the studies selected

			Stu	udy			
Variable	Bang	et al ^[16]	Paik	et al ^[15]	Park et al ^[14]		
	EUS	ERCP	EUS	ERCP	EUS	ERCP	
п	33	34	64	61	15	15	
Age (yr), mean (SD)	69.4 (12.6)	69.2 (11.6)	64.8 (12.5)	68.4 (10.5)	66.8 (8)	65.4 (9.3)	
Etiology of MBO	Pancreas (n = 33)	Pancreas (n = 31); pancreatic metastasis (n = 3)	Pancreas (<i>n</i> = 38); cholangiocarcinoma (<i>n</i> = 3); gallbladder (<i>n</i> = 4); papilla (<i>n</i> = 5); gastric (<i>n</i> = 4); duodenal (<i>n</i> = 2); other (<i>n</i> = 8)	Pancreas (<i>n</i> = 40); cholangiocarcinoma (<i>n</i> = 8); gallbladder (<i>n</i> = 4); papilla (<i>n</i> = 3); gastric (<i>n</i> = 2); duodenal (<i>n</i> = 1); hepatocellular carcinoma (<i>n</i> = 1); other (<i>n</i> = 2)	Pancreas (n = 14); cholangiocarcinoma (n = 1)	Pancreas (<i>n</i> = 13); metastatic lymph node (<i>n</i> = 2)	

EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography; MBO: Malignant biliary obstruction.

after ERCP failure^[11-13]. Moole *et al*^[22] recently published a systematic review and metaanalysis comparing PTBD and EUS-guided drainage as alternatives to be employed after failed ERCP, demonstrating that the latter was superior, as has been corroborated by other authors^[23].

Because of improvements in the technique and accessories over time, some authors have reported EUS-guided biliary drainage as a first-line modality in patients presenting with factors predictive of difficult biliary access by ERCP (*e.g.*, altered anatomy, duodenal obstruction, and previous duodenal stent)^[24-28]. Okuno *et al*^[27] published a prospective study of 20 patients undergoing EUS-guided hepaticogastrostomy with a 6-mm self-expanding metallic stent. The rates of technical success, clinical success, and adverse events were 100%, 95%, and 15%, respectively. In a recent multicenter cohort study^[28], EUS-guided biliary drainage was compared with ERCP in patients with an indwelling duodenal stent. The authors identified a trend toward higher technical and clinical success rates in the EUS group and found no difference regarding adverse events. Finally, Nakai *et al*^[29] published a retrospective study comparing primary and rescue EUS-guided biliary drainage in terms of the rates of technical success and adverse events, both of which the authors found to be similar between the two approaches.

In a recent retrospective study of patients with distal biliary obstruction, Kawabuto *et al*^[30] demonstrated that EUS-guided choledochoduodenostomy was similar to transpapillary stenting in terms of the rates of clinical success and adverse events, although the duration of the procedure was shorter and there were no cases of pancreatitis among the patients submitted to the former. Therefore, the EUS-guided procedure was considered a plausible first-line method to address MBO. Subsequent RCTs comparing those techniques have shed light on the matter^[14-16].

The availability of three high-quality RCTs allowed us to perform a consistent meta-analysis that will likely contribute to making daily practice more evidence based. Our analysis of technical success is extremely reliable because of the similar definitions employed and homogenous results among the three studies. However, the clinical success analysis lacked consistency because of indirectness due to different outcome definitions. In addition, Bang *et al*^[16] included cross-over procedures in the data report, which impeded the intention-to-treat analysis. Therefore, caution should be taken in drawing conclusions based on the results of this analysis.

During our evaluation of the duration of procedures, we found high heterogeneity among the studies. Such true heterogeneity is likely attributable to the participation of endoscopists with different levels of expertise. In addition, various stents have been used in biliary drainage. Paik *et al*^[15] employed insulated delivery systems to perform EUS-guided drainage, which probably shortened the duration of the procedure in their EUS group and promoted heterogeneity. Our analysis showed equivalence between the two methods regarding the duration of the procedure. It should be borne in mind that, whereas ERCP is a well-established technique, EUS-guided drainage is still in development, and its duration could therefore become shorter in the near future.

As to the safety of the procedure, our analysis showed similar rates of adverse events after EUS and ERCP. Although there was no difference between the two



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Table 2 Jadad	Table 2 Jadad scale scores for the studies selected											
Study.			ladad scale s	coring								
Study	Randomization	Appropriate randomization	Blinding	Appropriate blinding	Losses described	Total						
Bang et al ^[16]	Yes	Yes	No	No	Yes	3						
Paik et al ^[15]	Yes	Yes	No	No	Yes	3						
Park <i>et al</i> ^[14]	Yes	Yes	No	No	Yes	3						

approaches regarding the overall rates, there was a substantial difference regarding the types of adverse events observed. In the ERCP group, the most common complication was pancreatitis, which was not reported in the EUS group. Conversely, pneumoperitoneum and biliary peritonitis were reported only in the EUS group, although none of patients required surgical intervention. Cholangitis was reported in both groups: 7 cases in the ERCP group and 4 in the EUS group.

The stent patency was equivalent for both methods, although the largest study^[15] did not provide standard deviation values and was therefore excluded. That significantly reduced the size of the sample evaluated in the stent patency analysis.

Finally, the results of our analysis of the stent dysfunction rate favored EUS-guided drainage. That might be explained by the fact that this method allows a puncture far from the tumor rather than through it, thus avoiding tumor ingrowth or overgrowth. Although EUS-guided drainage can promote stent dysfunction due to food bolus impaction, that risk does not seem to outweigh its advantages. It should also be borne in mind that the employment of diverse stents for biliary drainage could be a confounding factor in the analysis of stent patency and dysfunction.

None of the RCTs evaluated in our systematic review described a cost-effectiveness analysis. Because the differences between ERCP and EUS-guided drainage are still slight, such information might create a tipping point to recommend one approach over the other. Future trials should address this knowledge gap.

Limitations

Our study has some limitations. First, a lack of standard deviation data precluded the inclusion of the largest trial in the stent patency analysis, thus limiting our ability to draw conclusions regarding that aspect. Second, different definitions of clinical success resulted in a very low quality of evidence, also precluding any firm conclusions. Finally, the small number of RCTs included constitutes a major limitation. Future studies might therefore contradict our results. Nevertheless, to our knowledge, this is the first meta-analysis comparing ERCP and EUS-guided biliary drainage in MBO. Our findings could have significant clinical implications for the management of patients with MBO.

Conclusion

In patients with distal MBO, EUS-guided drainage shows rates of technical success, clinical success, adverse events, and stent patency similar to those of ERCP. The rates of stent dysfunction appear to be lower for stents placed under EUS guidance. Cost-effectiveness studies might solidify the role of EUS-guided drainage in the management of MBO.



Table 3 Quality (certainty) of evidence of the studies selected, as determined by the GRADE criteria

Parameter		Inconsistence	lu dina atu a a a	luon no sision	Dublication bios	Overall containty of ovidence	
No. of patients (studies)	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Overall certainty of evidence	
Technical success							
222 (3 RCTs)	Not serious	Not serious	Not serious	Serious ^a	None	Moderate	
Clinical success							
155 (2 RCTs)	Serious ^b	Not serious	Serious ^c	Serious ^a	None	Very low	
Procedure duration							
222 (3 RCTs)	Not serious	Very serious ^d	Serious ^e	Serious ^a	None	Very low	
Adverse events							
222 (3 RCTs)	Not serious	Very serious ^d	Not serious	Serious ^a	None	Very low	
Stent patency							
97 (2 RCTs)	Serious ^b	Not serious	Serious ^e	Serious ^a	None	Very low	
Stent dysfunction							
155 (2 RCTs)	Not serious	Not serious	Serious ^e	Not serious	Strongly suspected	Low	

^aNo significant difference found. ^bIncomplete outcome data in one study. ^cStudies used different criteria for clinical success. ^d*I*² > 75%. ^eOutlier identified. RCTs: Randomized clinical trials.

	EU	S	ERC	CP .		Risk difference	Risk difference
Study or subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95%CI	M-H, Fixed, 95%CI
Bang 2018	30	33	32	34	30.2%	-0.03 [-0.16, 0.09]	
Palk 2018	60	64	55	61	56.3%	0.04 [-0.06, 0.13]	
Park 2018	13	15	14	15	13.5%	-0.07 [-0.28, 0.15]	
Total (95%CI)		112		110	100.0%	0.00 [-0.07, 0.07]	
Total events	103		101				
Heterogenelty: $Chi^2 = 3$	1.16, df = 2	(P = 0.5)	56); $I^2 = 0^6$	%			
Test for overall effect:	Z = 0.04 (P	= 0.97)					-0.2 -0.1 0 0.1 0.2 ERCP EUS

Figure 2 Forest plot of technical success. M-H: Mantel-Haenszel test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

	EU	S	ERC	CP		Risk difference	Risk difference
Study or subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95%CI	M-H, Fixed, 95%CI
Palk 2018	54	64	52	61	60.6%	-0.01 [-0.13, 0.12]	_
Park 2018	13	15	13	15	19.4%	0.00 [-0.24, 0.24]	
Total (95%CI)		79		76	100.0%	-0.01 [-0.12, 0.10]	
Total events	67		65			- / -	
Heterogenelty: $Chi^2 = 1$	0.00, df = 1	(P = 0.9)	$95); I^2 = 0^9$	%			
Test for overall effect:							-0.2 -0.1 0 0.1 0.2 ERCP EUS

Figure 3 Forest plot of clinical success. M-H: Mantel-Haenszel test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

		EUS		E	RCP			Mean difference	Mean difference
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95%CI	IV, Random, 95%CI
Bang 2018	24.2	9.2	33	22.4	13.5	34	37.9%	1.80 [-3.72, 7.32]	
Palk 2018	5	2.25	64	11	2.75	61	45.8%	-6.00 [-6.88, -5.12]	•
Park 2018	43	24	15	31	21	15	16.2%	12.00 [-4.14, 28.14]	
Total (95%CI)			112			110	100.0%	-0.12 [-8.20, 7.97]	
Heterogenelty: Tua ² Test for overall effect				f = 2 (<i>P</i>	= 0.00	2); <i>I</i> ² =	84%		-20 -10 0 10 20 EUS ERCP

Figure 4 Forest plot of procedure duration in minutes. IV: Inverse variance test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

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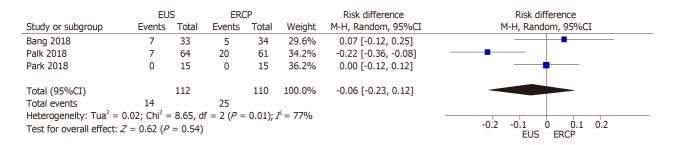


Figure 5 Forest plot of adverse events. M-H: Mantel-Haenszel test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

	E	EUS		EF	RCP			Mean difference	Mean difference
Study or subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95%CI	IV, Fixed, 95%CI
Bang 2018 Park 2018	182 2 379	1.25 55	33 15	170 403	37 84	34 15	92.6% 7.4%	12.00 [-2.40, 26.40] -24.00 [-74.81, 26.81]	
Total (95%CI) Heterogenelty: Chi ² = Test for overall effect	,	•		$I^2 = 449$	%	49	100.0%	9.32 [-4.53, 23.18]	-50 -25 0 25 50 ERCP EUS

Figure 6 Forest plot of stent patency. IV: Inverse variance test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

	EU	S	ERC	CP		Risk difference	Risk difference
Study or subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95%CI	M-H, Fixed, 95%CI
Bang 2018	1	33	1	34	0.0%	0.00 [-0.08, 0.08]	
Palk 2018	10	64	24	61	80.6%	-0.24 [-0.39, -0.09]	
Park 2018	2	15	4	15	19.4%	-0.13 [-0.42, 0.15]	
Total (95%CI)		79		76	100.0%	-0.22 [-0.35, -0.08]	
Total events	12		28				-
Heterogeneity: $Chi^2 = 0$	0.41, df = 1	(P = 0.5)	52); $I^2 = 0^6$	%			
Test for overall effect:							-0.2 0.1 0 0.1 0.2 EUS ERCP

Figure 7 Forest plot of stent dysfunction requiring intervention. M-H: Mantel-Haenszel test; EUS: Endoscopic ultrasound; ERCP: Endoscopic retrograde cholangiopancreatography.

ARTICLE HIGHLIGHTS

Research background

Endoscopic retrograde cholangiopancreatography (ERCP) is currently the gold standard palliation approach for distal malignant biliary obstruction (MBO) but as endoscopic ultrasound (EUS)-guided techniques develop and became more commonly available question arises whether EUS-guided biliary drainage cloud be a first line method for treatment of distal MBO.

Research motivation

EUS-guided biliary drainage and ERCP are recognized endoscopic approaches for palliation of MBO. Our initial motivation was to compare EUS and ERCP techniques for primary drainage of distal MBO. By performing a systematic review and meta-analysis following a rigorous methodological approach we aimed to increase the available knowledge regarding endoscopic palliation of MBO.

Research objectives

To perform a systematic review and meta-analysis comparing EUS and ERCP as primary methods of biliary drainage in distal MBO regarding technical success, clinical success, duration of the procedure, adverse events, stent patency and stent dysfunction.

Research methods

We conducted a systematic review and meta-analysis based on the PRISMA Statement and registered on PROSPERO international database. We searched the Medline, *Excerpta Medica*, and Cochrane Central Register of Controlled Trials databases. Only randomized clinical trials (RCTs) comparing EUS and ERCP for primary drainage of MBO were eligible. We assessed the risk of biases using the Jadad score and the quality of evidence using the Grading of Recommendations Assessment, Development and Evaluation criteria.

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Research results

Three RCTs were included in the final analysis comprising a total of 222 patients (112 submitted to EUS and 110 submitted to ERCP). The stent dysfunction rate was significantly lower in the EUS group (MD = -0.22%; 95% CI: -0.35, -0.08; P = 0.001; $l^2 = 0\%$). There were no statistically significant difference regarding technical success, clinical success, duration of the procedure, adverse events and stent patency among the compared techniques.

Research conclusions

In palliative drainage of distal MBO, EUS-guided and ERCP drainage presents similar rates of technical success, clinical success, adverse events, and stent patency. The rates of stent dysfunction appear to be lower for stents placed under EUS guidance.

Research perspectives

We considered meaningful to stablish a present evaluation of both techniques and as the procedures continue to develop, further widespread and new technologies emerge, we encourage that additional RCT's and meta-analisys are performed.Cost-effectiveness studies might solidify the role of EUS-guided drainage in the management of MBO.

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