Research Article

Effects Comparison between Endoscopic Papillary Large Balloon Dilatation and Endoscopic Sphincterotomy for Common Bile Duct Stone Removal

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Endoscopic sphincterotomy (EST) is a treatment of choice for stone extraction and is now most frequently used. The study was to compare the efficacy of endoscopic papillary large balloon dilatation (EPLBD) and endoscopic sphincterotomy (EST) for common bile duct stone removal. Trials comparing the effects between EPLBD and EST treatment were searched according to the study protocol. Overall stone removal rate, complete removal rate in 1st session, treatment duration, mechanical lithotripsy using rate, and overall complication rate were compared using risk ratio (RR) and mean difference (MD) and their 95% confidence interval (CI) via RevMan 5.2 software. For overall stone removal rate, two therapies showed similar effect, but EPLBD showed better overall stone removal rate for stone >10 mm in diameter. For complete stone removal rate in 1st session, no difference was found, even for those with stone >10 mm in diameter; EPLBD showed longer treatment duration, higher mechanical lithotripsy using rate obvious overall complications rate, and more serious bleeding, whereas there were no significant differences for perforation, hyperamylasemia, pancreatitis, and cholecystitis/cholangitis. EPLBD showed better efficacy in certain conditions compared to EST, however with shortcomings, such as more duration, higher mechanical lithotripsy using rate, and bleeding.

1. Introduction

Common bile duct stones are present in about 4–10 percent of patients who have undertaken cholecystectomy [1]. Reported incidence of common bile duct stone varies from 5% to 11% at the time of cholecystectomy [2, 3]. The vast majority of common bile duct stone mainly originates from gallbladder [4]. Its signs and symptoms are variable, ranging from being completely asymptomatic to complications, such as biliary colic, cholangitis, jaundice, or pancreatitis [5].

Endoscopic sphincterotomy (EST), first described in 1974, is a treatment of choice for stone extraction and is now most frequently used [6]. Although EST is relatively safe, it is reported to induce high level of overall complication rates, as well as short-term risks, including bleeding, perforation, and pancreatitis. Endoscopic papillary large balloon dilatation (EPLBD) had been introduced as an alternation for EST to manage the common bile duct stones, for its lower frequency in perforation and hemorrhage [7]. However, in another study comparing these two treatments, EST showed lower pancreatitis occurrence than EPLBD and 2 patients died of pancreatitis because of the EPLBD extraction [8]. It remained as a controversial issue whether EPLBD has a better effect than EST.

In this study, we conducted a meta-analysis to compare the data collected in trials published between 2004 and 2013. Overall stone removal rate, complete stone removal rate in 1st session, treatment duration, mechanical lithotripsy using rate, and overall/each complications rate were compared between the two therapies.

2. Methods

2.1. Information Sources and Searches. A search of the literature was conducted for studies that reported the EPLBD versus EST for common bile duct stone removal.

15, 16.5, or 18 mm

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		TABLE 1: Ch	aracteristics of the included 1	0 prospective studies.	
Study	Design*	Country	Bile duct stones	Intervention (<i>n</i>)	Dilated balloon catheter
Lin et al., 2004 [9]	D, S, R	China	Multiple, mean = 8 mm	EPLBD (51) or EST (54)	10–12 mm
Heo et al., 2007 [6]	D, S, R	Korea	Multiple, mean = 16.0–15.0	EST plus EPLBD (100) or EST (100)	12–20 mm
Itoi et al., 2009 [10]	S	Japan	Large	Small EST plus EPLBD (53) or EST (48)	10–20 mm
García-Cano et al., 2009 [11]	D, S	Spain	Multiple, mean = 3 mm	EST plus EPLBD (31) or EST (60)	10–20 mm
Kim et al., 2009 [12]	D, S, R	South Korea	Large, ≥15 mm	Small EST plus EPLBD (27) or EST (28)	15, 16.5, or 18 mm
Kim et al., 2011 [13]	S	Korea	Large, ≥10 mm	Small EST plus EPLBD (72) or EST (77)	12–20 mm
Stefanidis et al., 2011 [14]	D, S, R	Greece	Large, >12 mm	Full EST plus EPLBD (45) or EST plus ML (45)	10–20 mm
Oh and Kim, 2012 [15]	S, R	Korea	Large, >10 mm	EPLBD (40) or EST (43)	10-20 mm
Teoh et al., 2013 [16]	D, S, R	China	Large, >13 mm	Limit EST plus EPLBD (73) or EST (78)	15 mm

Multiple, mean = 12.7-13.2

*D: double blinded; S: single center; R: randomization.

S, D

China

Li et al., 2014 [17]

The PubMed, Web of Science, Medline, Science Citation Index, EMBASE, China National Knowledge Infrastructure, Wanfang Database, and China Biomedical Database were searched to identify double blinded research (D, S, R), single center trials (S), and random clinical trials (RCTs published) in the field of common bile duct stone removal between 2004 and 2013. The keywords used in literature searches included the following: bile duct stones, endoscopic sphincterotomy, and endoscopic papillary balloon dilatation.

2.2. Eligibility Criteria and Outcome Measure. The inclusion criteria were the following: (i) the included studies were designed to compare the therapeutic effects of endoscopic sphincterotomy and endoscopic papillary large balloon dilatation or the combination of the two therapeutics and (ii) the publications could be written in any language. Reports of duplicated studies were excluded by examining the author list, parent institution, sample size, and results.

The primary outcome was overall stone removal rate, overall/complete stone removal rate in 1st session, treatment duration, mechanical lithotripsy using rate, and overall/each complications rate.

2.3. Assessment of Study Quality. Two authors (Shan Lei and Chen Li) independently assessed the quality of the included studies according to the descriptions provided by the authors of the included trials. The methodological quality of the trials was assessed based on adequate sequence generation, allocation concealment, blinding, management of incomplete outcome data, and early stopping for benefit [18].

2.4. Study Selection and Data Collection. Two authors (Shan Lei and Chen Li) independently screened titles and abstracts for potential eligibility and the full texts for final eligibility. We extracted the data from the included trials independently

for quantitative analyses, and any disagreement was subsequently resolved by discussion. The quantitative data included the country, stone diameters, interventions, and dilated balloon catheter.

Small EST plus EPBD (232) or

EST (230)

2.5. Synthesis of Results. In this meta-analysis, we used a random effect model because of the anticipated variability among trials with regard to patient populations [19, 20]. The measure of association used in this meta-analysis was the risk ratio (RR) or mean difference (MD) with a 95% confidence interval (CI). The summary RR with the 95% CI was calculated by the RevMan 5.2 software using the random or fixed effect model. A statistically significant result was assumed when the 95% CI did not include one.

Heterogeneity was explored using a Chi-square test; P < 0.05 represents that there is heterogeneity of effect size. In addition, the quantity of heterogeneity was measured using the I^2 statistic, which calculates the percentage of total variation across studies caused by heterogeneity rather than sampling errors. $I^2 = 25\%$ was considered as low heterogeneity and 50% as moderate, while 75% was considered as high.

3. Results

3.1. Literature Search and Population. Fifteen trials were finally identified in the English literature. Two trials were removed after blind review because they did not compare the result of EST and EPLBD, 2 reviews were also excluded, and the data of 1 literature was also excluded for the information being incomplete. Characteristics of the final 10 prospective studies [6, 9–17] are shown in Table 1. Patient numbers with EST and EPLBD treatment were 763 and 769 (Table 1).

3.2. Comparison of the Overall Stone Removal Rate of EPLBD and EST Treatments. The rate ratios for overall stone removal

Study or subgroup	EPL Events	.BD Total	ES Events	ST Total	Weight	Risk ratio M-H, fixed, 95% CI	Risk ratio M-H, fixed, 95% CI
Lin et al., 2004	48	51	53	53	7.4%	0.94 [0.87, 1.02]	
Teoh et al., 2013	71	73	78	78	10.7%	0.97 [0.93, 10.2]	
Heo et al., 2007	97	100	98	100	13.8%	0.99 [0.95, 1.03]	
Kim et al., 2009	27	27	28	28	4.0%	1.00 [0.93, 1.07]	- + -
Itoi et al., 2009	53	53	47	48	7.0%	1.02 [0.97, 1.08]	+
Oh and Kim, 2012	39	40	41	43	5.6%	1.02 [0.94, 1.11]	
Li et al., 2014	222	232	215	230	30.5%	1.02 [0.98, 1.07]	-
Kim et al., 2011	70	72	73	77	10.0%	1.03 [0.96, 1.09]	
García-Cano et al., 2009	29	30	57	61	5.3%	1.03 [0.94, 1.14]	_ _
Stefanidis et al., 2011	44	45	41	45	5.8%	1.07 [0.97, 1.19]	+
Total (95% CI)		723		763	100.0%	1.01 [0.99, 1.03]	•
Total events	637		674				· · · · · · · · · · · · · · · · · · ·
Heterogeneity: $\chi^2 = 9.0$ Test for overall effect: <i>Z</i>			$I^2 = 1\%$				0.5 0.7 1 1.5 2 Favours EST Favours EPLBD

Ctur day ou ouch amoun	EPL	BD	ES	Т	Weight	Risk ratio			Risk ratio)	
Study or subgroup	Events	Total	Events	Total	weight	M-H, fixed, 95% CI		M-H	, fixed, 9	5% CI	
Kim et al., 2009	27	27	28	28	9.2%	1.00 [0.93, 1.07]			_		
Itoi et al., 2009	53	53	47	48	16.3%	1.02 [0.97, 1.08]					
Oh and Kim, 2012	39	40	41	43	12.9%	1.02 [0.94, 1.11]			_ 		
Kim et al., 2011	70	72	73	77	23.1%	1.03 [0.96, 1.09]					
Stefanidis et al., 2011	44	45	41	45	13.4%	1.07 [0.97, 1.19]			+		
Li et al., 2014	88	90	75	86	25.1%	1.12 [1.03, 1.22]				-	
Гotal (95% CI)		327		327	100.0%	1.05 [1.02, 1.09]			•		
Total events	321		305								
Heterogeneity: $\chi^2 = 6.4$	2, $df = 5 (P$	P = 0.27);	$I^2 = 22\%$				0.5	0.7	1	1.5	2
Test for overall effect: Z]	Favours EST	Fav	ours EPLI	BD

(a)

(b)

FIGURE 1: Rate ratio for overall stone removal rate of EPLBD and EST treatment (a) and overall stone removal rate for stone >10 mm in diameter (b).

rate for patients taking EPLBD therapies were similar to that of those taking EST therapies (RR: 1.01, 95% CI: 0.99-1.03; P = 0.35) (Figure 1(a)), but, for the patient with stone >10 mm in diameter, the rate ratios for overall stone removal rate of EPLBD were higher than those of EST (RR: 1.05, 95% CI: 1.02–1.09, P < 0.05 (Figure 1(b)). The result showed that EPLBD had better efficacy than EST method for stones larger than 10 mm in terms of overall removal rate. The rate ratio for complete stone removal rate in 1st session of EPLBD was similar to that of EST (RR: 1.07, 95% CI: 0.98-1.16, P = 0.11) (Figure 2(a)), and the rate ratio for complete stone >10 mm in diameter removal rate in 1st session for EPLBD and EST was also similar (RR: 1.11, 95% CI: 0.99-1.24, P = 0.08) (Figure 2(b)). Thus, there was no significant difference between the two kinds of treatment in terms of complete stone removal rate in 1st session. In this metaanalysis for complete stone >10 mm in diameter removal rate in 1st session, there was apparent heterogeneity (P = 0.01, $I^2 = 64\%$), so random effect model was used. What is more, for the other outcomes, fixed effect model was used for there was no apparent heterogeneity.

3.3. Comparison for Treatment Duration and Mechanical Lithotripsy Using Rate for EPLBD and EST. There was significant heterogeneity in the assessment of treatment duration $(P < 0.00001, I^2 = 90\%)$, so random effect model was used. It turned out that the two kinds of treatment were significantly different (MD = -5.05, 95% CI: -9.55~ -0.54, P = 0.03; Figure 3(a)), and EPLBD showed longer treatment duration. Fixed effect model was used for mechanical lithotripsy assessment for no heterogeneity ($P = 0.13, I^2 = 37\%$), and the RR value was 0.47 with 95% CI being between 0.37 and 0.60 (P < 0.00001; Figure 3(b)). EPLBD treatment had higher mechanical lithotripsy using rate compared with EST treatment.

3.4. Comparison for Complications Rate for EPLBD and EST. No apparent heterogeneities were detected between the studies in terms of overall complications rate (P = 0.22, $I^2 = 24\%$) and each complications rate (bleeding: P = 0.11, $I^2 = 41\%$; perforation: P = 0.63, $I^2 = 0\%$; hyperamylasemia: P = 0.22, $I^2 = 24\%$; pancreatitis: P = 0.98, $I^2 = 0\%$; cholecystitis/cholangitis: P = 0.64, $I^2 = 0\%$; then fixed

Study or subgroup	EPL	BD	ES	Т	Weight	Risk ratio	Risk ratio
Study of subgroup	Events	Total	Events	Total	weight	M-H, fixed, 95% CI	M-H, fixed, 95% CI
Heo et al., 2007	83	100	87	100	16.1%	0.95 [0.85, 1.07]	
Lin et al., 2004	44	51	47	53	13.7%	0.97 [0.84, 1.13]	
Teoh et al., 2013	65	73	69	78	16.4%	1.01 [0.90, 1.13]	_ _
Kim et al., 2009	23	27	23	28	8.3%	1.04 [0.82, 1.31]	
Itoi et al., 2009	51	53	41	48	15.1%	1.13 [0.99, 1.28]	⊢ ∎−−
Kim et al., 2011	63	72	57	77	12.7%	1.18 [1.01, 1.39]	
Li et al., 2014	200	232	162	230	17.7%	1.22 [1.11, 1.35]	
Total (95% CI)		608		614	100.0%	1.07 [0.98, 1.16]	
Total events	529		486				
Heterogeneity: $\tau^2 = 0$	$0.01; \chi^2 = 16$	5.45, df = 0	5(P = 0.01)); $I^2 = 6$	4%		0.5 0.7 1 1.5 2
Test for overall effect	Z = 1.59 (H	P = 0.11)					Favours EST Favours EPLBD

Study or subgroup	EPL	BD	ES	бТ	Weight	Risk ratio		Ris	sk ratio		-
Study of subgroup	Events	Total	Events	Total	weight	M-H, fixed, 95% CI		M-H, fix	ed, 95%	o CI	
Kim et al., 2009	23	27	23	28	21.6%	1.04 [0.82, 1.31]					
Kim et al., 2011	63	72	57	77	52.7%	1.18 [1.01, 1.39]					
Teoh et al., 2013	28	33	26	31	25.7%	1.01 [0.82, 1.25]				—	
Total (95% CI)		132		136	100.0%	1.11 [0.99, 1.24]				•	
Total events	114		106								
Heterogeneity: $\chi^2 =$	1.66, df = 2	(P = 0.44)	; $I^2 = 0\%$				0.5	0.7	1	1.5	2
Test for overall effect]	Favours EST]	Favours EPL	BD

(b)

FIGURE 2: Rate ratio for complete stone removal rate in 1st session of EPLBD and EST treatment (a) and complete stone removal rate in 1st session for stone >10 mm in diameter (b).

Study or subgroup	EPL Mean	BD SD	Total	ES Mean	SD	Total	Weight	Mean difference IV, random, 95% CI	Mean difference IV, random, 95% CI
Teoh et al., 2013 Li et al., 2014 Itoi et al., 2009 Lin et al., 2004	24.3 38.6 32 44	12.87 15.5 8 3	73 232 53 51	27.2 47.1 40 45	16.9 20.2 8 3	78 230 48 53	21.7% 24.9% 25.2% 28.2%	-2.90 [-7.67, 1.87] -8.50 [-11.79, -5.21] -8.00 [-11.12, -4.88] -1.00 [-2.15, 0.15]	
Total (95% CI)			4.9			409	100.0%	-5.05 [-9.55, -0.54]	
Heterogeneity: $\tau^2 =$ Test for overall effect				P < 0.000	001); I ²	= 90%			-10 -5 0 5 10 Favours EPLBD Favours EST

					(a)			
Study or subgroup	EPL Events	BD Total	ES Events	T Total	Weight	Risk ratio M-H, fixed, 95% CI	Risk ra M-H, fixed	
Itoi et al., 2009	3	53	12	48	7.4%	0.23 [0.07, 0.75]		
Li et al., 2014	28	232	80	230	47.3%	0.35 [0.23, 0.51]		
Kim et al., 2011	6	72	15	77	8.5%	0.43 [0.18, 1.04]		
Oh and Kim, 2012	4	40	9	43	5.1%	0.48 [0.16, 1.43]		
Lin et al., 2004	1	51	2	53	1.2%	0.52 [0.05, 5.56]		
Teoh et al., 2013	21	73	36	78	20.5%	0.62 [0.40, 0.96]		
Heo et al., 2007	7	100	8	100	4.7%	0.88 [0.33, 2.32]		
Kim et al., 2009	9	27	9	28	5.2%	1.04 [0.49, 2.21]		_
Total (95% CI)		648		657	100.0%	0.47 [0.37, 0.60]	•	
Total events	79		171					
Heterogeneity: $\chi^2 = 1$	1.16, df = 7 (P = 0.13);	$I^2 = 37\%$				0.15 0.2 1	5 20
Test for overall effect:	Z = 6.23 (P + 1)	< 0.00001))				Favours EPLBD	Favours EST

(b)

FIGURE 3: Comparison of treatment duration (a) and mechanical lithotripsy (b) using rate of EPLBD and EST treatment.

Study or subgroup	EPL	BD	ES	σT	Weight	Risk ratio	Risk ratio
Study of subgroup	Events	Total	Events	Total	weight	M-H, fixed, 95% CI	M-H, fixed, 95% CI
Stefanidis et al., 2011	2	45	9	45	7.3%	0.22 [0.05, 0.97]	
Lin et al., 2004	4	51	18	53	14.3%	0.23 [0.08, 0.64]	
Teoh et al., 2013	8	73	18	78	14.1%	0.47 [0.22, 1.02]	
Kim et al., 2011	9	72	19	77	14.9%	0.51 [0.25, 1.05]	
Li et al., 2014	19	232	32	230	26.0%	0.59 [0.34, 1.01]	
Itoi et al., 2009	2	53	3	48	2.5%	0.60 [0.11, 3.46]	
Heo et al., 2007	5	100	7	100	5.7%	0.71 [0.23, 2.18]	_
García-Cano et al., 2009	2	30	5	61	2.7%	0.81 [0.17, 3.95]	
Oh and Kim, 2012	10	40	11	43	8.6%	0.98 [0.47, 2.06]	_ _
Kim et al., 2009	8	27	5	28	4.0%	1.66 [0.62, 4.44]	+
Total (95% CI)		723		763	100.0%	0.57 [0.44, 0.75]	•
Total events	69		127				·
Heterogeneity: $\chi^2 = 11.86$	6, df = 9 (P)	= 0.22);	$I^2 = 24\%$				0.01 0.1 1 10 100
Test for overall effect: $Z =$	4.02 (P <	0.00001)					Favours EPLBD Favours EST

					(a)		
Study or subgroup	EPI Events	LBD Total	EST Events	Г Total	Weight	Risk ratio M-H, fixed, 95% CI	Risk ratio M-H, fixed, 95% CI
Bleeding							· · ·
Lin et al., 2004	1	51	14	53	28.4%	0.07 [0.01, 0.54]	
Heo et al., 2007	0	100	2	100	5.2%	0.20 [0.01, 4.11]	
Li et al., 2014	4	232	12	230	24.9%	0.33 [0.11, 1.01]	
Oh and Kim, 2012	4	40	7	43	14.0%	0.61 [0.19, 1.94]	
Teoh et al., 2013	8	73	10	78	20.0%	0.85 [0.36, 2.05]	
Stefanidis et al., 2011	1	45	1	45	2.1%	1.00 [0.06, 15.50]	
Kim et al., 2009	4	27	2	28	4.1%	0.07 [0.41, 10.41]	
García-Cano et al., 2009	2	30	1	61	1.4%	4.07 [0.38, 43.08]	
Subtotal (95% CI)		598		638	100.0%	0.53 [0.34, 0.84]	
Total events Heterogeneity: $\chi^2 = 11.85$,	$df = \frac{24}{7}(P =$	$= 0.11); I^2 =$	49 41%				•
Test for overall effect: $Z =$	2.69 (P = 0	.007)					
Perforation							
Teoh et al., 2013	0	73	2	78	41.3%	0.21 [0.01, 4.37]	
Stefanidis et al., 2011	0	45	1	45	25.6%	0.33 [0.01, 7.97]	
Kim et al., 2011	0	72	1	77	24.8%	0.36 [0.01, 8.60]	
Oh and Kim, 2012	1	40	0	43	8.2%	3.22 [0.13, 76.82]	
Subtotal (95% CI)		230		243	100.0%	0.53 [0.13, 2.08]	-
Hyperamylasemia							
Kim et al., 2011	3	72	9	77	55.9%	0.36 [0.10, 1.26]	
Lin et al., 2004	3	51	4	53	25.2%	0.78 [0.18, 3.31]	
Kim et al., 2009	4	27	3	28	18.9%	1.38 [0.34, 5.61]	_
Subtotal (95% CI)		150		158	100.0%	0.66 [0.31, 1.40]	◆
Total events Heterogeneity: $\chi^2 = 2.03$, o Test for overall effect: $Z =$	10 df = 2 (P = 1.08 (P = 0	0.36); $I^2 =$.28)	16 2%				
O and ama addition							
		30	3 9	61	6.2%	0.29 [0.02, 5.36]	
García-Cano et al., 2009	0			77	23.0%	0.59 [0.21, 1.69]	
García-Cano et al., 2009 Kim et al., 2011	5	72	2	70			
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013	5 2	73	3	78	7.7%	0.71 [0.12, 4.14]	
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012	5 2 2	73 40	3 3	43	7.6%	$\begin{array}{c} 0.71 \\ [0.12, 4.14] \\ 0.72 \\ [0.13, 4.07] \end{array}$	
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014	5 2 2 13	73 40 232	3 3 17	43 230	7.6% 45.1%	$\begin{array}{c} 0.71 \\ 0.12, 4.14 \\ 0.72 \\ 0.73, 4.07 \\ 0.76 \\ 0.38, 1.52 \end{array}$	
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012	5 2 2	73 40	3 3	43	7.6%	$\begin{array}{c} 0.71 \\ [0.12, 4.14] \\ 0.72 \\ [0.13, 4.07] \end{array}$	
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014 Heo et al., 2007 Subtotal (95% CI) Total events	5 2 13 4 26	73 40 232 100 547	3 3 17 4 39	43 230 100	7.6% 45.1% 10.6%	0.71 [0.12, 4.14] 0.72 [0.13, 4.07] 0.76 [0.38, 1.52] 1.00 [0.26, 3.89]	•
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014 Heo et al., 2007 Subtotal (95% CI)	5 2 13 4 26	73 40 232 100 547	3 3 17 4 39	43 230 100	7.6% 45.1% 10.6%	0.71 [0.12, 4.14] 0.72 [0.13, 4.07] 0.76 [0.38, 1.52] 1.00 [0.26, 3.89]	•
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014 Heo et al., 2007 Subtotal (95% CI) Total events Heterogeneity: $\chi^2 = 0.76$, 6 Test for overall effect: $Z =$ <i>Cholecystitis/cholangitis</i>	5 2 13 4 26	73 40 232 100 547 $(0.98); I^2 = 0.16$	3 3 17 4 39 0%	43 230 100	7.6% 45.1% 10.6%	0.71 [0.12, 4.14] 0.72 [0.13, 4.07] 0.76 [0.38, 1.52] 1.00 [0.26, 3.89]	•
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014 Heo et al., 2007 Subtotal (95% CI) Total events Heterogeneity: $\chi^2 = 0.76$, 6 Test for overall effect: $Z =$ <i>Cholecystitis/cholangitis</i> Kim et al., 2009	5 2 13 4 26	73 40 232 100 547	3 3 17 4 39 0%	43 230 100 589 28	7.6% 45.1% 10.6%	0.71 [0.12, 4.14] 0.72 [0.13, 4.07] 0.76 [0.38, 1.52] 1.00 [0.26, 3.89]	•
García-Cano et al., 2009 Kim et al., 2011 Teoh et al., 2013 Oh and Kim, 2012 Li et al., 2014 Heo et al., 2007 Subtotal (95% CI) Total events Heterogeneity: $\chi^2 = 0.76$, ζ Test for overall effect: $Z = Cholecystitis/cholangitis$ Kim et al., 2009 Lin et al., 2004	5 2 2 13 4 4 4 6 f = 5 (P = 0) 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0	$\begin{array}{c} 73\\ 40\\ 232\\ 100\\ 547\\ \hline \\ (0.98); I^2 = \\ .16)\\ 27\\ 51\end{array}$	3 3 17 4 39 0%	43 230 100 589 28 53	7.6% 45.1% 10.6%	0.71 [0.12, 4.14] 0.72 [0.13, 4.07] 0.76 [0.38, 1.52] 1.00 [0.26, 3.89] 0.71 [0.44, 1.15]	•
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(b)

FIGURE 4: Comparison of overall complications rate (a) and each complications rate (b) of EPLBD and EST treatment.

effect model was used for the meta-analyses. EPLBD showed obvious overall complications rate (RR = 0.57, 95% CI = 0.44~ 0.75, P < 0.0001; Figure 4(a)) and more serious bleeding (RR = 0.53, 95% CI = 0.34~0.84, P = 0.007; Figure 4(b)) than EST, while there were no significant differences between these two treatments for perforation (RR = 0.36, 95% CI = 0.13~2.08, P = 0.36), hyperamylasemia (RR = 0.66, 95% CI = 0.31~1.40, P = 0.28), pancreatitis (RR = 0.71, 95% CI = 0.44~1.15, P = 0.16), and cholecystitis/cholangitis (RR = 0.62, 95% CI = 0.28~1.37, P = 0.24) (Figure 4(b)).

4. Discussion

EST used to be the most commonly used treatment for the removal of bile duct stones. Nevertheless, it is commonly reported that EST could induce substantial procedure associated risks, such as increased incidence of ascending cholangitis [6]. Endoscopic papillary balloon dilatation uses a small balloon catheter, while EPLBD uses a larger balloon (>12 mm) after the mid-incision EST to remove large common bile duct stones. EPLBD would theoretically combine the advantages of balloon dilation and sphincterotomy by increasing stone extraction efficacy while minimizing complications of them. We performed this meta-analysis to compare the effect of EST and EPLBD for common bile duct stones. Even no significant difference existed in the overall stone removal rate, but, for patients with stones >10 mm in diameter, EPLBD could give better overall stone removal rate; however, this treatment needs more duration and will induce higher mechanical lithotripsy using rate with more serious overall complications rate and bleeding, compared with EST. The result was consistent with the study of Fujita et al. [21]. Nevertheless, many studies have recently proposed that EPLBD will not cause complications if performed under strictly established guidelines [22]. Thus, it is of great importance to follow the guidelines during the treatment of EPLBD.

We reviewed previous meta-analysis and found that patients received EPLBD therapy which was associated with shorter length of hospitalization, whereas there was no significance for clearance rates and morbidity/mortality [4]. However, these conclusions may be unreliable, because only 2 included trials reported these outcomes. The current study is based on 10 prospective studies, which are potentially more robust than previous meta-analysis.

There are few limitations of our current study: (1) not all the included literatures studied all the assessment parameters, such as complete stone removal rate in 1st session and treatment duration; (2) the trials adapted different design, including D, S, R, and R; (3) we did not conduct sensitive analysis to test the reliability of the results.

In conclusion, the EPLBD treatment needs more treatment duration and mechanical lithotripsy using rate compared to EST, but, for patients with stones >10 mm in diameter, EPLBD could give better overall stone removal rate. What is more, EPLBD would produce obvious overall complications rate and bleeding, compared with EST.

Conflict of Interests

The authors declare that there is no conflict of interests regarding the publication of this paper.

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