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## Robotic versus conventional laparoscopic surgery for rectal cancer: systematic review and meta-analysis

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**Purpose:** Robotic surgery (RS) overcomes the limitations of previous conventional laparoscopic surgery (CLS). Although meta-analyses have been published recently, our study evaluated the latest comparative surgical, urologic, and sexual results for rectal cancer and compares RS with CLS in patients with rectal cancer only.

Methods: We searched three foreign databases (Ovid-MEDLINE, Ovid-Embase, and Cochrane Library) and five Korean databases (KoreaMed, KMbase, KISS, RISS, and KisTi) during July 2013. The Cochrane Risk of Bias and the Methodological Index for Non-Randomized were utilized to evaluate quality of study. Dichotomous variables were pooled using the risk ratio (RR), and continuous variables were pooled using the mean difference (MD). All meta-analyses were conducted with Review Manager, V. 5.3.

**Results:** Seventeen studies involving 2,224 patients were included. RS was associated with a lower rate of intraoperative conversion than that of CLS (RR, 0.28; 95% confidence interval [CI], 0.15–0.54). Time to first flatus was short (MD, –0.13; 95% CI, –0.25 to –0.01). Operating time was longer for RS than that for CLS (MD, 49.97; 95% CI, 20.43–79.52,  $I^2 = 97\%$ ). International Prostate Symptom Score scores at 3 months better RS than CLS (MD, –2.90; 95% CI, –5.31 to –0.48,  $I^2 = 0\%$ ). International Index of Erectile Function scores showed better improvement at 3 months (MD, –2.82; 95% CI, –4.78 to –0.87,  $I^2 = 37\%$ ) and 6 months (MD, –2.15; 95% CI, –4.08 to –0.22,  $I^2 = 0\%$ ).

**Conclusion:** RS appears to be an effective alternative to CLS with a lower conversion rate to open surgery, a shorter time to first flatus and better recovery in voiding and sexual function. RS could enhance postoperative recovery in patients with rectal cancer.

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Key Words: Robotic surgical procedures, Rectal neoplasms

#### **INTRODUCTION**

Laparoscopic colorectal surgery has been proven to be as safe and effective as open surgery and offers various advantages [1]. Conventional laparoscopic surgery (CLS) provides good magnification and illumination for colorectal surgeons, which improve visualization of pelvic structures [2]. CLS for patients with colorectal diseases results in better short-term morbidity than that from open surgery [3]. Although CLS has revolutionized surgical management of colorectal cancers over the past two decades, it has several technical limitations, particularly when excising rectal cancer, including a limited range of instrument motion in the narrow pelvic cavity, related loss of dexterity, and an inadequate visual field associated with an unstable camera view [1,4]. Robotic systems have been widely adopted for urological, gynecological, and rectal surgery. In particular, a robotic system is favorable for deep and narrow spaces, such as the pelvis or mediastinum [5,6].

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Since the first robot-assisted colectomy was performed in 2001 [7], a number of studies have been published on the use of robotic systems in colorectal surgery. In particular, robotic surgery (RS) overcomes the limitations of previous CLS, including a motion filter for tremor-free surgery, high definition three-dimensional imaging, an easily controlled camera on a stable platform, and increased space for the operating instruments [8]. Currently, Baik et al. [9] reported the feasibility and safety of robot-assisted tumor-specific mesorectal excision of rectal cancer in terms of oncologic outcomes. Also, Bae et al. [10] reported robotic pelvic lymph node dissection for rectal cancer was feasible and safe. A few systematic reviews and meta-analyses comparing outcomes between RS and CLS have been conducted [11-14]. However, there is some limitations of the studies because they analyzed safety and efficacy but included noncomparative studies or mixed colonic and rectal cancers.

Therefore, we evaluated the safety and efficacy of RS by performing a systematic review and meta-analysis focusing on rectal cancer studies that compared RS and CLS.

#### **METHODS**

#### Search strategy and study selection

We searched three foreign databases (Ovid-MEDLINE, Ovid-Embase, and Cochrane Library) and five Korean databases (KoreaMed, KMbase, KISS, RISS, and KisTi) on July 23, 2013. We searched pertinent MeSH or EMTREE terms (rectal neoplasms, rectal cancer, and rectal carcinoma or adenocarcinoma or tumor), intervention terms (robotics, computer-assisted surgery, telerobot, remote operation, remote surgery, and Da Vinci) and their combinations. No language was restricted, and no filters were applied. The references from published reviews were also assessed to seek additional publications. A total of 1,664 records were ultimately identified through the search. After removing duplicates, 1,302 studies were imported for the first title and abstract screening.

A total of 54 articles were obtained to conduct the full-text review, and 23 articles were identified to be potentially relevant. We excluded 9 of 23 publications because 4 may have had overlapping cohorts, and 5 studies did not report appropriate results. We considered sample size, publication year, study design, and the number of outcomes of interest to choose highquality data with a low risk of bias for the publications with possible overlap. Finally, we add 3 articles by hand searching. Total seventeen publications were ultimately selected for the meta-analysis (Fig. 1).

#### Inclusion and exclusion criteria

Two authors (S.L. and J.H.K.) independently reviewed all abstracts. We retrieved full-text copies of all studies that

potentially met the inclusion criteria based on a review of the abstract. If both authors agreed that a study did not meet the eligibility criteria, we excluded the study. If we disagreed, we resolved the conflict by discussion and consensus or by consulting a third review team member. The inclusion criteria were: (1) compared outcomes of RS and CLS in patients with rectal cancer and (2) if multiple studies were reported for the same patient population, either the one of higher quality or the most recent study was used. Studies fulfilling the following criteria were excluded: (1) the outcomes and parameters of patients with rectal cancer were not reported clearly; (2) it was impossible to extract appropriate data from the published results; (3) the studies were not original articles and included gray literature, such as letters, editorials, and expert opinions or reviews without original data; and (4) abstracts, theses, casereports, and studies lacking a control group.

#### Data extraction and quality assessment

Each author extracted the data independently from each study and compared them, resolving disagreement by discussion: (1) first author and year of publication; (2) study design; (3) country in which the study was performed; (4) number of subjects operated on with each technique; (5) study population demographic and clinical characteristics; (6) clinical outcomes, conversion rates, time to oral diet, time to bowel function recovery, estimated blood loss (EBL), bowel obstruction, complications (anastomotic leakage, urinary retention, intrapelvic abscess, or intrapelvic bleeding); (7) oncologic outcomes (number of lymph nodes extracted); and (8) urological and sexual function (e.g., International Prostate Symptom Score [IPSS] and International Index of Erectile Function [IIEF]).

Two authors (S.L. and J.H.K.) assessed all of the studies included for methodological quality according to the Risk of Bias for Randomized Controlled Trials (RCTs) [15] and the Methodological Index for Non-randomized Studies (MINORS) for nonrandomized comparative studies [16]. The information is summarized in Table 1 and Fig. 2.

#### **Statistical analysis**

Dichotomous variables were pooled using the risk ratio (RR), and continuous variables were pooled using the mean difference (MD). RR and MD were calculated and reported with 95% confidence intervals (CIs). Statistical heterogeneity among the studies was analyzed using the chi-square and Q-tests [17]. A fixed effects model was used for studies with low or moderate statistical heterogeneity ( $I^2 < 25$ , 25%–50%), and a random effects model was used for studies with high statistical heterogeneity ( $I^2 > 50\%$ ). Meta-analyses of dichotomous variables were performed using the Mantel-Haenszel (M-H) method, and continuous variables were analysed using the

0	Voor		Study	Stud	ły size	(u)	Sex,	M:F	Age (	(yr)	BMI (k	g/m²)	CR	Ē	Tumor size	e (cm)	Robotic	(e 4:10).
	IEdi	COUNTRY	type	Total	RS	CLS	RS	CLS	RS	CLS	RS	CLS	RS (	CLS	RS	CLS	technique <sup>V</sup>	quiry
[2]	2008	Korea	RCT	36	18	18	14:4	14:4	$57.3 \pm 6.3$	$62.0 \pm 9.0$	$22.8 \pm 1.8$	$24.0 \pm 2.5$	,	Т	I	I	R-TME	ı
[24]	2009	Italy	RCT	99	29	37	11:18	12:25	$68 \pm 10.0$	$69 \pm 10.0$	$24 \pm 6.2$	$25.4 \pm 6.44$	$\sim$	2	ı	ı	R-LAR	ı
_	2010	Italy	PCS	50	25	25	18:7	17:8	69 (33–83)	62 (42–77)	24.6 (18.5–31.2)	26.5 (18.1–35.0)	ı.	ı	I	I	R-TME	14
al.	2010	Romania	RCS	122	38	84	23:15	51:33	53 ± 11.3	60 ± 12.3	ı	ı	ı.	ī	ı	ı	R-TME	10
21]	2011	NSA	PCS	82	41	41	25:16	25:16	63.5 (48–87)	63.7 (42–88)	25.7 (17.1–34.0)	26.7 (16.8–40.3)	33	18	ı	ı	R-TME	13
[6]	2011	Korea	RCS	140	52	88	28:24	70:53	57.3 ± 12.3	65.1 ± 10.3	23.7 ± 2.4	$23.6 \pm 3.3$	12	10	ı	ı	R-proctec- tomy R-TME	13
30]	2011	USA	RCS	60	30	30	19:11	19:11	$53.9 \pm 11.0$	$56.3 \pm 12.2$	$28.2 \pm 4.2$	$27.0 \pm 4.3$	4	4	ı	ı		12
[28]	2012	Korea	RCS	304	154	150	105:49	109:41	59.1 ± 12.2	62.3 ± 10.9	23.4 ± 3.1	$23.1 \pm 3.0$	35	12	ı		R-TME R-ISR R-LAR R-APR	
33]	2012	Korea	PCS	69	30	39	18:12	20:19	$54.13 \pm 8.5$	$56.85 \pm 11.1$	24.36 ± 2.4	$24.01 \pm 2.2$	19	15	ŀ	ı	R-TME	21
	2012	Korea	RCS	29	17	12	,	I	I	I	ı	I	ı.	ı	ı	I	R-proctec- tomy	10
	2013	Italy	RCS	100	50	50	30:20	30:20	66.0 ± 12.1	65.7 ± 11.6	ı	ı	34	28	I	I	R-TME	12
ıl.	2013	Turkey	RCS	64	27	37	ı	I	I	I	28.3 (19.3–30.8)	26.75 (19–40)	4	8	4 (0–15) 3	(6-0) 9.9	R- LAR	10
[18]	2013	Korea	RCS	330	165	165	104:61	97:68	61.2 ± 11.4	$60.4 \pm 11.8$	23.1 ± 2.8	$23.2 \pm 3.1$	24	23 3	.4±1.8 3	.3 ± 2.2	R-proctec- tomy	14
2]	2013	Korea	RCS	80	40	40	28:12	25:15	$57.3 \pm 12.1$	$63.6 \pm 10.6$	$23.9 \pm 2.4$	$24.3 \pm 3.1$	32	20 2	$.8 \pm 1.9$ 3	$.2 \pm 1.9$	R-ISP	11
et al.	2013	USA	RCS	72	13	59	13:0	57:2	$67.9 \pm 2.1$	64.9 ± 1.2		I	10	32	ı	I	R-APR R-LAR	12
32]	2014	Taiwan	RCS	64	36	28	21:15	17:11	55.9 (30-89)	54.9 (25-88)	23.78	23.32	28	28	I	ı	R-ISR	12
27]	2015	Korea	RCS	556	278	278	182:96	184:94	$57.4 \pm 11.6$	$58.3 \pm 10.4$	$23.5 \pm 2.9$	$23.7 \pm 3.3$	91	79 2	.9±1.9 3	i.1 ± 2.1	R-TME	16
	.																	

Table 1. Characteristics and quality assessment of the eligible studies

Values are presented as mean ± standard deviation or median (range) unless otherwise indicated. RS, robotic surgery; CLS, conventional laparoscopic surgery; BMI, body mass index; CRT, chemoradiotherapy; RCT, randomized controlled clinical trial; PCS, prospective cohort study; RCS, retrospective cohort study; R-TME, robotic rectal anterior resection with total mesorectal excision; R-ISP, robot-assisted intersphinteric resection; R-LAR, robotic low anterior resection; R-APR, robotic abdominoperitoneal resection.

<sup>al</sup>Quality assessment was done using the Methodological Index for Non-randomized (MINORS).

#### Annals of Surgical Treatment and Research 2015;89(4):190-201





**Fig. 1.** Flow diagram of the study selection process. RS, robotic surgery; CLS, conventional laparoscopic surgery.

inverse variance method. We conducted all meta-analyses with Review Manager, V. 5.3 (RevMan, Copenhagen, Denmark: The Nordic Cochrane Centre, The Cochrane Collaboration, 2014) and used the two-tailed test of significance (P < 0.05).

#### **Ethical consideration**

This study was reviewed and approved by the Institutional Review Board at the National Evidence-Based Healthcare Collaborating Agency (NECA): (approval number: NECA IRB 13-013)

#### RESULTS

#### **Study characteristics**

Table 1 shows the characteristics of the 17 studies and their patients. The 17 studies included 2,224 patients: 1,043 in the RS group and 1,181 in the CLS group. Two studies were RCTs, four were prospective cohort studies, and eleven were retrospective cohort studies. Of the 17 studies, 8 were conducted in Korea, 3 were conducted in Italy, 3 in the USA, 1 in Turkey, 1 in Taiwan, and 1 in Romania. All studies were published during the last 7 years (2008–2015). The risk of bias in the RCT studies was high (Fig. 2). The quality of the cohort studies, except Kim (2012), was poor (Table 1).

#### **Conversion rates**

Ten studies [18-27] reported a conversion rate to open



Fig. 2. The risk of bias for eligible randomized controlled trials.

surgery. In all, 8 patients (1.19%) undergoing RS and 50 (6.50%) undergoing CLS required conversion to open surgery (RR, 0.28; 95% CI, 0.15–0.54), with no heterogeneity among studies (P = 0.98,  $I^2 = 0$ %) (Table 2, Fig. 3).

#### Time to oral diet

Time to liquid diet was reported in six studies [2,18,22,27-29] comparing RS and CLS. The pooled estimates using a fixed

O. the second seco	No. of	Study si	ize (n)	Heterog	geneity	Effect	Statistical	Analysis	Pooled effect	
Outcome	study	RS	CLS	P-value	l <sup>2</sup> (%)	measure	method	model	RR (95% CI)	P-value
Clinical outcomes										
Conversion rates	10	674	769	0.980	0	RR	H-M	FE	0.28 (0.15 to 0.54)	<0.001
Time to oral diet	9	727	805	0.160	36	MD	2	FE	-0.18 (-0.36 to 0.00)	0.050
Time to recovery of bowel function	8	762	850	0.710	0	MD	2	FE	-0.13 (-0.25 to -0.01)	0.030
Estimated blood loss	8	751	796	0.001	71	MD	2	RE	-9.03 (-33.11 to 15.05)	0.460
Operating time	10	821	844	<0.001	97	MD	2	RE	49.97 (20.43 to 79.52)	<0.001
Duration of hospitalization	10	821	917	0.006	61	MD	2	RE	-0.53 (-1.26 to 0.21)	0.160
Complication										
Anastomotic leakage	10	819	925	0.950	0	RR	H-M	ΕE	0.87 (0.65 to 1.17)	0.360
Urinary retention	Ŋ	299	389	0.770	0	RR	H-M	FE	0.87 (0.47 to 1.61)	0.650
Intrapelvic bleeding	9	680	676	0.710	0	RR	H-M	ΕE	1.23 (0.51 to 2.94)	0.640
Intrapelvic abscess	Ŋ	566	602	0.950	0	RR	H-M	FE	1.21 (0.57 to 2.60)	0.620
Wound infection	7	625	717	0.950	0	RR	H-M	Ħ	0.76 (0.32 to 1.81)	0.540
Oncological outcomes										
No. of lymph nodes extracted	8	647	732	0.060	48	MD	2	Ħ	0.20 (-0.54 to 0.94)	0.600
Urological function										
IPSS (3 mo after surgery)	2	44	54	0.590	0	MD	2	FE	-2.90 (-5.31 to -0.48)	0.020
IPSS (6 mo after surgery)	2	44	54	0.490	0	MD	2	FE	-0.45 (-2.58 to 1.68)	0.680
IPSS (12 mo after surgery)	2	09	69	0.490	0	MD	2	Ħ	-0.90 (-1.93 to 0.14)	0.090
Sexual function <sup>a)</sup>										
IIEF (3 mo after surgery)	2	32	32	0.210	37	MD	2	Ħ	-2.82 (-4.78 to -0.87)	0.005
IIEF (6 mo after surgery)	2	32	29	0.930	0	MD	2	FE	-2.15 (-4.08 to -0.22)	0.030
RS, robotic surgery; CLS, conventional la effects; RE, random effects; IPSS, Internatio <sup>a</sup> Change from baseline after surgery.	paroscopic mal Prostate	surgery; Cl, Symptom Sc	confidence :ore; IIEF, In	interval; RR, ternational Inc	risk ratio; N dex of Erecti	AD, mean di le Function.	fference; M-F	l, Mantel-Ha	ienszel; IV, inverse varianc	e; FE, fixed

Table 2. Clinical and oncological outcomes

Annals of Surgical Treatment and Research 2015;89(4):190-201

**ASTR** 

	RS	6	CL	S		Risk ratio		Risk ratio
Study or subgroup	Events	Total	Events	Total	Weight	M-H, Fixed, 95% CI	Year	M-H, Fixed, 95% CI
Baik SH et al [25]	0	18	2	18	5.9%	0.20 [0.01, 3.89]	2008	
Patriti A et al [24]	0	29	7	37	15.7%	0.08 [0.01, 1.42]	2009 –	
Popescu I et al [22]	2	38	9	84	13.3%	0.49 [0.11, 2.17]	2010	
Bianchi PP et al [23]	0	25	1	25	3.6%	0.33 [0.01, 7.81]	2010	
Baek JH et al [21]	3	41	9	41	21.4%	0.33 [0.10, 1.14]	2011	
Shin JY [20]	0	17	1	12	4.1%	0.24 [0.01, 5.45]	2012	
Fernandez et al [26]	1	13	10	59	8.6%	0.45 [0.06, 3.24]	2013	
Kang J et al [18]	1	165	3	165	7.1%	0.33 [0.04, 3.17]	2013	
D'Annibale A et al [19]	0	50	6	50	15.4%	0.08 [0.00, 1.33]	2013 —	
Cho et al [27]	1	278	2	278	4.8%	0.50 [0.05, 5.48]	2015	
Total (95% CI)		674		769	100.0%	0.28 [0.15, 0.54]		•
Total events	8		50	~				•
Heterogeneity: Chi <sup>2</sup> = 2	2.64, df =	= 9 (P =	= 0.98); I	<sup>2</sup> = 0%	)			
Test for overall effect:	Z = 3.86	(P = 0	.0001)				0.002	0.1 1 10 500
								Favours RS Favours CLS

Fig. 3. Forest plot and meta-analysis of conversion to open surgery. RS, robotic surgery; CLS, conventional laparoscopic surgery; M-H, Mantel-Haenszel; Cl, confidence interval; df, degree of freedom.



Fig. 4. Forest plot and meta-analysis of time to first flatus. RS, robotic surgery; CLS, conventional laparoscopic surgery; SD, standard deviation; CI, confidence interval; df, degree of freedom.

effects model showed no significant difference between RS and CLS (MD, -0.18 days; 95% CI, -0.36 to 0.00), with moderate heterogeneity (P = 0.16, I<sup>2</sup> = 36%) (Table 2).

#### Time to bowel function recovery

Eight studies [2,18,20,22,25,27-29] analyzed time to first flatus. The pooled estimates using a fixed effects model showed a significant difference between RS and CLS (MD, -0.13 days, 95% CI, -0.25 to -0.01), with no heterogeneity among studies (P = 0.71, I<sup>2</sup> = 0%) (Table 2, Fig. 4).

#### **Estimated blood loss**

EBL was reported in eight studies [2,18,20,22,24,27,28,30] comparing RS and CLS. The pooled estimates using a random effects model showed no significant difference between RS and CLS (MD, -9.03 mL; 95% CI, -33.11 to 15.05), with heterogeneity (P = 0.001, I<sup>2</sup> = 71%) (Table 2).

#### **Operating time**

Operating time was reported in ten studies [2,18,20,22,24,25,

27-30] comparing RS and CLS. The pooled estimates using a random effects model showed significantly longer operating time for RS than that for CLS (MD, 49.97 minutes; 95% CI, 20.43–79.52). However, this result had high heterogeneity (P < 0.001,  $I^2 = 97\%$ ) (Table 2, Fig. 5).

#### **Duration of hospitalization**

Hospitalization duration was reported in ten studies [2,18,20,22,24,25,27-30] comparing RS and CLS. The pooled estimates using a random effects model showed no significant difference between RS and CLS (MD, -0.53 days; 95% CI, -1.26 to 0.21), with heterogeneity (P = 0.006, I<sup>2</sup> = 61%) (Table 2).

#### Complications

#### Anastomotic leakage

Anastomotic leakage was reported in ten studies [2,18,19,22-24,26-28,31] comparing RS and CLS. No significant difference in bowel anastomotic leaks was observed between RS and CLS (RR, 0.87; 95% CI, 0.65–1.17), with no heterogeneity (P = 0.95,  $I^2 = 0\%$ ) (Table 2).

#### Annals of Surgical Treatment and Research 2015;89(4):190-201



Study or subgroup	Mean	RS SD	Total	Mean	CLS SD	Total	Weight	Mean difference IV, Fixed, 95% Cl	Year		Mear IV, Fix	n diffe ked, §	erence 95% CI		
Baik SH et al [25]	217.1	51.6	18	204.3	51.9	18	9.3%	12.80 [-21.01. 46.61]	2008						
Patriti A et al [24]	202	12	29	208	7	37	10.6%	-6.00 [-10.92, -1.08]	2009			-			
Popescu I et al [22]	212	47.23	38	182	37.23	84	10.3%	30.00 [13.00, 47.00]	2010						
Park JS et al [29]	232.6	52.4	52	158.1	49.2	30	10.0%	74.50 [51.85, 97.15]	2011						
Patel CB et al [30]	237	56.8	30	181.6	52.5	30	9.7%	55.40 [27.72, 83.08]	2011					_	
Shin JY [20]	396.5	41.7	17	298.8	36.6	12	9.7%	97.70 [69.03, 126.37]	2012				-		
Baek SJ et al [28]	285.2	69.1	154	219.7	71.2	150	10.3%	65.50 [49.72, 81.28]	2012					_	
Park SY et al [2]	235.5	57.5	40	185.4	72.8	40	9.7%	50.10 [21.35, 78.85]	2013					-	
Kang J et al [18]	309.7	115.2	165	277.8	81.9	165	10.1%	31.90 [10.33, 53.47]	2013			-			
Cho et al [27]	361.6	91.9	278	272.4	83.8	278	10.4%	89.20 [74.58, 103.82]	2015						
Total (95% CI)			821			844	100.0%	49.97[20.43, 79.52]						•	
Heterogeneity: Tau <sup>2</sup>	= 2137	.54: Ch	i <sup>2</sup> = 28	5.80, d	f = 9 (P	< 0.0	0001); I <sup>2</sup>	= 97%	_				-		
Test for overall effec	:t: Z = 3	.32 (P =	= 0.00	09)	``		,,			-100	-50	0	50	100	C
				,						Fav	ours R	S Fa	avours (	CLS	

Fig. 5. Forest plot and meta-analysis of operation time. RS, robotic surgery; CLS, conventional laparoscopic surgery; SD, standard deviation; CI, confidence interval; df, degree of freedom.



Fig. 6. Forest plot and meta-analysis of International Prostate Symptom Score 3 months after surgery. RS, robotic surgery; CLS, conventional laparoscopic surgery; SD, standard deviation; CI, confidence interval; df, degree of freedom.

#### Urinary retention

Urinary retention was reported in five studies [2,18,24,26,29] comparing RS and CLS. The pooled estimates using a fixed effects model showed no significant difference between RS and CLS (RR, 0.87; 95% CI, 0.47–1.61), with no heterogeneity (P = 0.77,  $I^2 = 0\%$ ) (Table 2).

#### Intrapelvic bleeding

Intrapelvic bleeding was reported in six studies [18,24,25, 27,28,32] comparing RS and CLS. The pooled estimates using a fixed effects model showed no significant difference between RS and CLS (RR, 1.23: 95% CI, 0.51–2.94), with no heterogeneity (P = 0.71,  $I^2 = 0\%$ ) (Table 2).

#### Intrapelvic abscess

Intrapelvic abscess was reported in five studies [18,21,27, 29,30] comparing RS and CLS. The pooled estimates using a fixed effects model showed no significant difference between RS and CLS (RR, 1.21; 95% CI, 0.57–2.60), with no heterogeneity (P = 0.95,  $I^2 = 0\%$ ) (Table 2).

#### Wound infection

Wound infection was reported in seven studies [2,18,22,23,27, 29,31] comparing RS and CLS. The pooled estimates using a

fixed effects model showed no significant difference between RS and CLS (RR, 0.76; 95% CI, 0.32–1.81), with no heterogeneity (P = 0.95,  $I^2 = 0\%$ ) (Table 2).

#### Number of lymph nodes extracted

Eight studies [18-20,22,24,25,27,29] reported the numbers of lymph nodes extracted. No significant differences were observed in the total numbers of lymph nodes extracted between patients undergoing RS and CLS (MD, 0.20; 95% CI, -0.54 to 0.94), with no heterogeneity (P = 0.06, I<sup>2</sup> = 48%) (Table 2).

#### **Urological function (IPSS)**

Urological function was reported in three studies [2.19,33] comparing RS and CLS. The outcome was evaluated using IPSS with high scores corresponding with greater dysfunction. At three months follow-up, the pooled estimates using a fixed effects model showed significant difference of IPSS scores between RS and CLS (MD, -2.90; 95% CI, -5.31 to -0.48), with no heterogeneity (P = 0.59, I<sup>2</sup> = 0%) (Table 2, Fig. 6). At six month three months follow-up, the pooled estimates showed a no significant difference of IPSS scores between RS and CLS (MD, -0.45; 95% CI, -2.58 to 1.68), with no heterogeneity (P = 0.49, I<sup>2</sup> = 0%) (Table 2). Lastly, at the 12 months, there was no significant difference of IPSS score between RS and CLS (MD, -0.49, I<sup>2</sup> = 0%) (Table 2).

#### Seon Heui Lee, et al: Robotic surgery for rectal cancer

Study or subgroup	Mean	RS SD	Total	Mean	CLS SD	Total	Weight	Mean difference IV, Fixed, 95% CI	Year	Mean difference IV, Fixed, 95% Cl
Kim JY et al [33] Park SY et al [2]	8 3.8	5.8 3	18 14	13 5.9	6 3.2	17 15	25.0% 75.0%	-5.00 [-8.91, -1.09] -2.10 [-4.36, 0.16]	2012 2013	
Total (95% CI) Heterogeneity: Chi <sup>2</sup> Test for overall effe	<sup>2</sup> = 1.58, ct: Z = 2	df = ^ 2.83 (F	32 1 (P = 0 P = 0.0	0.21); I <sup>2</sup> 05)	= 379	32 %	100.0%	-2.82 [-4.78, -0.87]	-	-10 -5 0 5 10 Favours RS Favours CLS

Fig. 7. Forest plot and meta-analysis of International Index of Erectile Function score change from baseline to 3 months after surgery. RS, robotic surgery; CLS, conventional laparoscopic surgery; SD, standard deviation; CI, confidence interval; df, degree of freedom.

Study or subgroup	Mean	RS SD	Total	Mean	CLS SD	Total	Weight	Mean difference IV, Fixed, 95% CI	Year	ľ	Mean di √, Fixed,	fference , 95% Cl	
Kim JY et al [33] Park SY et al [2]	4.5 2.4	5 2.8	18 14	6.8 4.5	6 3.3	14 15	24.5% 75.5%	-2.30 [-6.20, 1.60] -2.10 [-4.32, 0.12]	2012 2013				
Total (95% CI) Heterogeneity: Chi <sup>*</sup> Test for overall effe	<sup>2</sup> = 0.01, ct: Z = 2	df = <sup>-</sup> 2.18 (F	32 1 (P = 1 P = 0.0	0.93); I <sup>2</sup> 3)	= 0%	29	100.0%	-2.15 [-4.08, -0.22]			5 0 JITS RS I	5 Favours C	+ 10 CLS

Fig. 8. Forest plot and meta-analysis of International Index of Erectile Function score change from baseline to 6 months after surgery. RS, robotic surgery; CLS, conventional laparoscopic surgery; SD, standard deviation; CI, confidence interval; df, degree of freedom.

-0.87; 95% CI, -1.93 to 0.19), with no heterogeneity (P = 0.52, I<sup>2</sup> = 0%) (Table 2).

#### Sexual function (IIEF)

Sexual function was reported in two studies [2,33] comparing RS and CLS. The outcome was evaluated using IIEF with high scores related to better sexual function. In this study, only erectile function was analyzed because Park et al. [2] reported assessment of male erectile function. At three months follow-up, the pooled estimates using a fixed effects model showed significant difference of IIEF scores between RS and CLS (MD, -2.82; 95% CI, -4.78 to -0.87), with little heterogeneity (P = 0.21, I<sup>2</sup> = 37%) (Table 2, Fig. 7). At six month three months follow-up, the pooled estimates showed a significant difference of IIEF scores between RS and CLS (MD, -2.15; 95% CI, -4.08 to -0.22), with no heterogeneity (P = 0.93, I<sup>2</sup> = 0%) (Table 2, Fig. 8).

#### **Publication bias**

A funnel plot analysis of the studies was performed in a meta-analysis reporting on overall postoperative complications after RS compared to those after CLS. None of the studies lay outside the 95% CI, and no evidence of publication bias or heterogeneity was detected among the studies except for operation time (Fig. 9).

#### DISCUSSION

There is currently moderate quality evidence that CLS for

total mesorectal excision (TME) has similar effects as those of open TME on long-term survival outcomes for treating rectal cancer [34]. Nevertheless, CLS remains a difficult procedure to perform due to inherent limitations, including the use of straight rigid instruments within a small working area, limited freedom, the fulcrum effect, and poor ergonomics [1,35]. In addition, CLS has been slow to gain popularity for treating rectal cancer, as a result of the technical difficulties associated with extensively dissecting the pelvis [36]. The da Vinci Surgical System (Intuitive Surgical, Sunnyvale, CA, USA) has been popularized for performing RS because it offers technical features that help overcome such difficulties. Adopting RS has facilitated the application of minimally invasive surgery to complex surgical operations in colorectal oncology. RS for rectal cancer is a technically feasible, safe procedure and can be performed with low morbidity and a low conversion rate, even for cases with advanced rectal cancer requiring complicated, robot-assisted, lateral lymph node dissection [37].

We conducted a systematic review and meta-analysis of comparative studies published during 2008–2015 to compare safety and effectiveness between RS and CLS in patients with rectal cancer. Although meta-analyses have been published recently [11.12.38.39], our study includes the most publications and reflects the latest comparative surgical results for rectal cancer. We also performed this meta-analysis to compare RS with CLS in patients with malignant diseases of the rectum only. Our results show two benefits of RS over CLS, and showed favorable urological and sexual function in RS group. Annals of Surgical Treatment and Research 2015;89(4):190-201

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Fig. 9. Funnel plot of the studies used in the meta-analysis. IPSS, International Prostate Symptom Score; IIEF, International Index of Erectile Function; SE, standard error; MD, mean difference; RR, risk ratio.

We found that the RS conversion rate for rectal cancer was significantly lower than that for CLS, indicating that RS provides better management of complex procedures and allowing more patients to benefit from minimally invasive surgery [39]. The reasons for conversion to open surgery included obesity with heavy mesentery, inability to identify important vascular structures, vascular injury, adhesions, and narrow pelvis; technical difficulties included stapler misfiring, inappropriate robotic arm placement, and robot malfunction [40]. The lower RS conversion rate may have been due to superior exposure and visualization of the operating field in the pelvis, thanks to the ability of the fixed fourth arm to grip and maneuver organs and the ability of the surgeon to move the three-dimensional camera as required [23]. Because conversion rate is related to a technical difficulty, surgical experience, and an intraoperative complication, it is good marker of the effectiveness of a robotic procedure [39]. The conversion rate for laparoscopic rectal cancer surgery is usually high, as 82 of 242 patients (34%) with rectal cancer were converted into open surgery in the MRC CLASICC trial [1]. The most common reasons were excess tumor fixation or uncertain tumor clearance, obesity, anatomical uncertainties, and tumor inaccessibility. Conversion to open surgery increases cost [41], and the patient no longer benefits from a laparoscopic approach. Conversion results in increased blood loss, prolonged postoperative hospital stay, and higher rates of intraoperative and postoperative complications. Moreover, conversion leads to higher postoperative mortality [42] and leads to poorer survival than those of laparoscopic or open rectal surgery [43]. Therefore, the lower conversion rate is clinically significant both in the short-term and long-term.

Another major advantage of RS was the marginally significant reduction in time to first flatus. The pooled data of our study show that time to first flatus was 0.13 days shorter following RS than that after LAS. The minimally invasive characteristics and technological advantages of a robotic system may have influenced this result. Baik et al. [25] explained that the da Vinci Surgical System robotic arms are used for retraction and dissection during the TME procedure, which reduces unnecessary procedures and minimizes iatrogenic tissue injury during retraction. These differences may have been the reason for the shorter time to first flatus in RS patients, but a more advanced study is needed before such a conclusion can be drawn. However, there is little clinical significance in these small differences.

Our meta-analysis demonstrated no difference in the number of retrieved lymph nodes. TME is widely accepted as the gold standard for RS and is one of the most important factors for reducing local recurrence and to assess the oncological adequacy of resection, Further studies should compare several secondary endpoints, such as margin clearance and mesorectal fascia integrity [44].

Our study showed favorable urological and sexual function in RS group. Specifically, IPSS scores 3 months after RS were better than those of the CLS and IIEF scores change from baseline to 6 months after RS were better than CLS. Damage to hypogastric nerves or sacral splanchnic nerves affect the patient's urological and sexual functions [45,46]. Also, when the CLS is used, it is more difficult to achieve autonomic nerve preservation than RS due to insufficient tension on the dissected planes [47,48]. Three-dimensional view conducted by the robotic system allowed the surgeon to better preserve the pelvic autonomic nerves. Besides, the use of the articulated mono-polar cautery hook allowed the surgeon to obtain better control of energy delivery, avoiding inopportune cauterization of the nervous bundle [19]. However, our results included a relatively small number of patients compared in both groups that the results need to be interpreted carefully.

This study had several important limitations. First, most studies in our review reported an insufficient follow-up period so many of the data were unsuitable to evaluate oncological outcomes, such as recurrence, metastasis, and mortality. Second, we could not determine whether the surgeon's learning curve affected the end result, as the studies included did not report comparisons between initial and subsequent RS experience.

In conclusion, RS provides an enhanced benefit to patients with rectal cancer in terms of postoperative recovery and the quality of the surgery. However, the quality of the evidence assembled does not support strong conclusions about most of the parameters of interest. In addition, RS is associated with a high cost and longer operating times. Therefore, findings supporting the safety and efficacy of RS need to be interpreted with caution because of the lack of RCTs. Only RCTs with a long-term follow-up could clearly determine whether the technological advantages of a robotic surgical system translate into favorable surgical and oncological outcomes. Further studies are required to evaluate the functional results and oncological outcomes associated with RS, such as the multicenter RCT of robot-assisted vs. laparoscopic resection for rectal cancer (ROLARR trial) [49].

#### **CONFLICTS OF INTEREST**

No potential conflict of interest relevant to this article was reported.

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