#### ORIGINAL RESEARCH

# Arm port vs chest port: a systematic review and meta-analysis

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**Background:** Two prevailing, totally implantable venous access ports are routinely utilized in oncology: chest port or arm port. This systematic review with meta-analysis was conducted to compare safety and efficiency of the two techniques.

**Methods:** We performed evidence acquisition intensively from PubMed, Embase, and Cochrane Library. Available comparative studies that evaluated both techniques were identified. The outcomes of interest included total complication events, procedure-related infections, thrombosis, intra-operative complications, mechanical complications, conversion rate, early port removal, and operating time.

**Results:** Thirteen comparative studies including 3,896 patients (2,176 for chest ports, and 1,720 for arm ports) were identified. The present study showed that arm port was associated with higher procedure conversion rate (2.51% in chest port group and 8.32% in arm port group; odd ratios [OR] 0.27, 95% confidence interval [CI] 0.15-0.46; p<0.001), but lower incidence of intra-operative complications (1.38% in chest port group and 0.41% in arm port group; OR 2.38, 95% CI 1.07–5.29; p=0.03). There were no between-group differences with respect to total complication events, procedure-related infections, thrombosis, mechanical complications, early port removal, and operating time. Subgroup analysis of patients under 60 years revealed that no significant difference was detected in intra-operative events (1.19% in chest port group and 0.02% in arm port group, OR 2.59, 95% CI 0.74–9.08; p<0.14), indicating that age may be a risk factor for intra-operative events. Sensitivity analysis did not change conclusions of all endpoints of interest.

**Conclusion:** Arm port is associated with higher procedure conversion rate, but lower incidence of intra-operative complications, and age may be a risk factor for intra-operative events.

**Keywords:** chest port, arm port, total implantable venous access port, systematic review, meta-analysis

# Introduction

Totally implantable venous access ports (TIVAPs), a long-term indwelling infusion system, was first introduced by Niederhuber in 1982.<sup>1</sup> With the increasing use of chemotherapeutic drugs in oncologic patients, TIVAP gains its popularity due to the fact that this system enables long-term administration of chemotherapeutic agents, attenuates the burden of chemotherapy, and thus greatly improves quality of life in oncologic patients, meanwhile, this system preserves peripheral vessels and prevents venous-related infections.<sup>2–4</sup> Two prevailing approaches are routinely utilized in oncology: chest ports or arm ports. Chest ports are most frequently implanted under the guidence of ultrasonography, via internal jugular vein, external jungular vein or subclavian vein puncture, while arm ports are placed through forearm veins such as basilic vein, cephalic vein or brachial vein, either through percutaneous

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© 2019 Li et al. This work is published and licensed by Dove Medical Press Limited. The full terms of this license are available at https://www.dovepress.com/terms.php you hereby accept the Terms. Non-commercial uses of the work are permitted without any further permission form Dove Medical Press Limited, provided the work is properly attributed. For permission for commercial use of this work, please see paragraphs 4.2 and 5 of our Terms (http://www.dovepress.com/terms.php). puncture or surgical cut-down, both generally showing satisfactory results with respect to technical success and low incidence of postprocedural complications.<sup>5,6</sup>

For years, the chest has been the most popular and reliable access site of implantation for long-term indwelling ports, however, chest ports might not be eligible for patients undergoing radiotherapy with chest wall involvement, or those with chest wall skin lesions, meanwhile, many female patients experience perception of "foreign body", "bra inconvenience", and discomfort when wearing safety belt or bag strap. In 1993, an alternative arm port procedure was introduced by surgeons at the School of Medicine, Yale University.<sup>7</sup> The arm ports have some unique advantages compared with chest ports, such as no risk for hemothorax or pneumothorax, lower incidence of arterial puncture,<sup>8</sup> better cosmetic outcome,<sup>9</sup> and more "bra convenience".<sup>10</sup> These advantages are especially beneficial to breast cancer patients requiring radiotherapy, flap transferring for reconstructive surgeries, as well as those patients with radioderrespiratory function.<sup>11</sup> matitis or compromised Nevertheless, the most significant disadvantage of arm port implantation is high incidence of failure (technically unsuccessful port placement), which is around 6.4%.<sup>12,13</sup>

Despite the wide application of TIVAPs, there is no consensus whether one or the other access site of implantation for TIVAP is clinically superior. Although several studies that compared chest access and arm access had been reported by various institutions, some of the results are conflicting and within small population of patients.<sup>14–16</sup> In the present study, we attempted to investigate and assemble the most comprehensive clinical data currently available in the literature to address a debatable issue: which approach, the arm port or the chest port, is more clinically beneficial to oncologic patients for the implantation of TIVAPs?

# Evidence acquisition

#### Clinical data search strategy

The present study was carried out according to the Metaanalysis Of Observational Studies in Epidemiology (MOOSE) recommendations and the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) protocol.<sup>17,18</sup> Systematic review registration was not available. Literature search was executed in August 2018, regardless of article types, regions or language of publication. Eligible studies were primarily from on-line databases of PubMed, Embase, and the Cochrane Library. Medical Subject Headings (MeSH) terms, as well as their combinations were manually searched via (Title/Abstract) as follows: basilic/cephalic/brachial/subclavian/jugular AND totally implantable venous access port/totally implantable venous access device/venous port/indwelling port, with "Related Articles" function applied to expand the search results. Corresponding author was contacted if additional information was required. When the same group of authors published multiple but similar reports of studies, the most comprehensive or the most recent result was taken into consideration.

# Inclusion and exclusion criteria

Inclusion criteria: 1) patients: patients who were diagnosed with malignancies requiring long-term indwelling ports, or those with gastrointestinal disorders requiring long-term venous port for parenteral nutrition support; 2) intervention: arm ports (basilic/cephalic/brachial vein insertion) vs chest ports (subclavian or jugular vein insertion), regardless of implantation methods, including surgical cut-down (peripheral vein cut-down method through surgical approach) and percutaneous puncture (radiology-guided placement, ultrasoundguided implantation, fluoroscopy-guided insertion, direct puncture, and so forth); 3) study types: all randomized controlled trials (RCTs), comparative studies available in the literature including cohort and case control studies which compared arm ports and chest ports were included; 4) outcomes: studies quantitatively reporting at least one of the outcomes described in the next section of this paper were included.

Exclusion criteria: 1) letters to the editor, editorials, reviews, case reports, laboratory-based animal studies, non-clinical studies, conference abstracts, and metaanalysis papers; 2) studies that failed to clearly clarify the outcomes of interest were excluded. If multiple access sites of implantation were compared in a single study (cephalic, jugular and subclavian vein), results of chest port/arm port category were combined.

# Data extraction

Two independent authors (YZ, GL) conducted data extraction. Any discrepancy was resolved by discussion and consultation with senior authors in this paper (HM and JZ). Data on clinical characteristics (age, diagnosis), type of port, and technique used (approach of insertion) were collected. Port-related complications were classified into four categories: intra-operative complications, infectious complications, thrombotic complications, and mechanical complications.<sup>19,20</sup>

The primary outcomes of interest were the incidences of port-related complications from implantation to removal of

TIVAP. Intra-operative complications included arterial puncture, hemothorax, and pneumothorax. Procedure-related infections, composed of port-site infection and catheterrelated bloodstream infection, were diagnosed in accordance with the latest guideline of the Infectious Diseases Society of America.<sup>21</sup> Procedure-related thrombotic complications were identified as any mural thrombi formed from the port catheter to the vessel lumen, giving rise to catheter occlusion, regardless of the presence of symptoms.<sup>22</sup> On the basis of Clavien-Dindo classification of surgical complications,<sup>23</sup> mechanical complications included port misfunctioning (such as deficiency of infusion or catheter occlusion), catheter fracture (defined as physical damage to the catheter of the port system), port dislocation (also named catheter migration or malposition), pinch-off syndrome, and hemorrhage. The second outcomes of interest were conversion to other access site of placement, early port removal (port removal before completion of treatment), and operating duration.

#### Quality assessment

Evaluation of RCT quality was subject to the Cochrane Collaboration's tool, through which risk of bias was assessed based on Cochrane Handbook for Systematic Reviews of Interventions.<sup>24</sup> Judgment of overall risk of bias for each included RCT was assessed as: low, moderate, or high.<sup>24</sup> RCTs with low risk of bias were regarded as high quality. Quality of observational study was assessed according to the Newcastle-Ottawa scale (NOS), consisting of three aspects: patient selection, comparability of the study groups, and assessment of outcome.<sup>25</sup> A quality score  $\geq$ 8 points was regarded as high quality study.

#### Statistical analysis

Review Manager 5.0 (Cochrane Collaboration, Oxford, UK) was used for this systematic review with metaanalysis. Weighted mean difference (WMD) was used to compare continuous variables, while odd ratio (OR) was employed to compare dichotomous variables. Ninety-five percent confidence interval (CI) was applied to all data throughout this study. When confronting with continuous variable expressed as mean and range value, standard deviation (SD) was converted as suggested by Hozo et al.<sup>26</sup>

Chi-squared test was used for the evaluation of statistical heterogeneity between studies, and significance level was set at p=0.10. For quantitative assessment of heterogeneity,  $I^2$  statistic was used, and high heterogeneity level was set at 75%.<sup>27</sup> The random-effects model was applied if between-study heterogeneity was high, in order to make more conservative estimates,<sup>28</sup> otherwise, the fixed-effects model was employed.<sup>24</sup>

Subgroup analyses were carried out for the outcomes of intra-operative complications, thrombotic complications, infectious complications, and mechanical complications. Available data were stratified based on patient's age, method of placement (puncture or surgical cut-down), whether antibiotic or thromboprophylaxis was applied, so as to investigate whether these factors influenced the outcomes. Sensitivity analysis was performed and only outcomes with at least two studies were included. We performed sensitivity analysis for high quality studies (RCT with low risk of bias and non-randomized studies with quality score  $\geq$ 8 points). We also used funnel plots to screen potential publication bias. A two-tailed *p*-value less than 0.05 was considered as statistical significance, except otherwise specified.

#### Results

A total of 914 possibly eligible studies were identified from databases (Figure 1). Thirteen studies including 3,896 cases (2,176 for chest ports, and 1,720 for arm ports) fulfilled the inclusion standard and were selected for final evidence synthesis. Agreement with respect to study selection between the two reviewers was 92%.

#### Characteristics of included studies

Table 1 depicts the characteristics of included studies in this study. Of these included studies, two<sup>29,30</sup> were RCTs, and



Figure I Flow diagram of study selection process.

Study	Year	Country	Design	Patients					Approach	Approach	Antibiotic	Coagulation	Matching
				Age, years (mean/ median)	Range	CP	AP	TIVAP	for CP	for AP	Prophylaxis	Prophylaxis	
Alahyane et al <sup>32</sup>	2010	Morroco	RC	43	16–76	340	240	NR	NR	NR	R	Yes	NR
Akahane et al <sup>33</sup>	2011	Japan	RC	64	2091	47	115	P-U Celsite	Puncture/	Puncture/	None	NR	1,2,4,5,7,8,9
									ultrasound	radiological			
Biffi et al <sup>29</sup>	2009	Italy	RCT	52	18–75	270	133	Bard port	Puncture/	Surgical cut-	NR	None	1,2,4,5,6,7
									ultrasound	down			
D'Angelo et al <sup>30</sup>	2002	Italy	RCT	61.5	17–75	25	25	NR	Puncture/	Surgical cut-	NR	NR	1,2,4,7
									ultrasound	down			
Goltz et al <sup>34</sup>	2012	Germany	RC	58.8	18–88	52	152	PowerPort	Puncture/	Puncture/	Yes	NR	1,2,4,5,7,8
									radiological	radiological			
Goltz et al <sup>10</sup>	2013	Germany	Ъ	55.8	19–84	25	25	PowerPort/P.	Puncture/	Puncture/	NR	NR	1,2,4,5,7
								A.S Port	radiological	radiological			
Kuriakose et al <sup>35</sup>	2002	USA	RC	58	1488	273	149	BardPort/	Puncture/	Puncture/	NR	Yes	1,2,4,5,6,7,9
								Meditec-	NR	radiological			
								R-Port					
Li et al <sup>36</sup>	2016	People's	RC	53.6	NR	237	107	Bard port	Puncture/	Puncture/	R	NR	1,2,4,5,7
		Republic of							direct	ultrasound			
		China											
Marcy et al <sup>37</sup>	2005	France	RC	55.7	RR	8	8	Bard/	Surgical cut-	Puncture/	None	NR	1,2,4,5,7,8
								BraunMedical	down	radiological			
Marcy et al <sup>38</sup>	2008	France	RC	59	20–83	112	113	Bard/	Surgical cut-	Puncture/	None	None	1,2,4,5,7,8,9
								BraunMedical	down	radiological			
Matiotti-Neto et al <sup>39</sup>	2017	USA	RC	NR	NR	247	195	NR	Puncture/	Surgical cut-	NR	NR	1,2,3,4,5,6,7
									fluoroscopy	down			
lorio et al <sup>31</sup>	2018	Italy	PC	59.5	NR	901	601	NR	Surgical cut-	Surgical cut-	Yes	NR	1,2,4,5,6,7,8
									down	down			
Shiono et al <sup>40</sup>	2014	Japan	RC	62.9	NR	342	257	Bard X-Port/	Puncture/	Puncture/	Yes	NR	1,2,4,5,7,8
								SlimPort	fluoroscopy	fluoroscopy			

two<sup>10,31</sup> were prospective non-randomized cohort studies. The remaining studies<sup>32-40</sup> were retrospective. As for indications, the majority of included studies were about cancer patients with upcoming chemotherapies, and most of the reported studies used commercial venous port devices (BardPort, PowerPort, P-U Celsite, BraunMedical, and Meditec-R-Port). For port implantation, the majority of studies used percutaneous puncture approach, while there were three studies<sup>31,37,38</sup> and four studies<sup>29,31,39</sup> which used direct surgical cut-down methods to implant chest port devices and arm port devices, respectively.

Based on Cochrane Handbook for Systematic Reviews of Interventions,<sup>24</sup> Table 2 summarizes the risk of bias of two included RCTs, one RCT was considered as moderate risk, while the other p< was regarded as low risk. In Table 3, risk of bias analysis of the remaining included studies was summarized in line with the NOS.<sup>25</sup> Among these non-randomized studies, two<sup>32,39</sup> had a quality score below seven, and the remaining studies were considered as high-quality.

#### Primary outcomes

The pooled data from 13 studies<sup>10,29-40</sup> that evaluated primary outcomes in 3,896 patients (Figure 2) revealed no significant differences for total complication events between the chest port and arm port groups (11.8% and 12.8%; OR 0.94, 95% CI 0.77-1.15; p=0.53). Eleven studies in which 3,266 patients were included thoroughly reported the incidences of intra-operative complications, thrombotic complications, infectious complications, and mechanical complications. For procedure-related infections, no statistically significant differences were observed between the chest port and arm port groups (3.15% and 3.37%; OR 1.11, 95% CI 0.74-1.66; p=0.63). Differences for procedure-related thrombotic complications and mechanical complications were also absent between the two groups (2.87% and 3.91%; OR 0.75, 95% CI 0.49–1.13; p=0.17; and 4.86% and 2.96%; OR 1.34, 95% CI 0.92-1.95; p=0.13, respectively). However, when comparing the difference for intra-operative complications, the incidence in chest port group was significantly higher (1.38% and 0.41%; OR 2.38, 95% CI 1.07–5.29; p=0.03) (Figure 3).

#### Secondary outcomes

Data on procedure conversion rate were obtained in six studies,<sup>26,29,31,38–40</sup> which assessed 1,314 and 914 patients in chest port group and arm port group, respectively. The procedure conversion rate was significantly higher in arm port group than that in chest port group (2.51% in chest port group and 8.32% in arm port group; OR 0.27, 95% CI 0.15-0.46; p < 0.001) (Figure 4A). When incidences of early port removal were analyzed, there were three studies<sup>29,36,39</sup> including 1,189 patients evaluated, and no significant difference was present between groups (7.29% and 4.37%; OR 1.27, 95% CI 0.26-6.19; p=0.77) (Figure 4A). As for operating time, three studies including 707 patients were reported, however, one study<sup>30</sup> presented operating time as "mean  $\pm$  standard deviation", one study<sup>39</sup> presented as "mean and range", and the last one<sup>31</sup> displayed as "mean and interquartile range". Using Hozo's<sup>26</sup> method, ranges were converted to estimated SD for further comparison.<sup>41</sup> Result of the analysis of operating time revealed that there were no statistically significant differences between chest port group and arm port group (WMD -4.31; 95% CI -17.81-9.19; p=0.53), with high between-study heterogeneity ( $l^2=0.98$ , p<0.01) (Figure 4B).

#### Subgroup analysis

In subgroup analysis of antibiotic prophylaxis, there were no statistically significant differences with respect to infection rate in comparison of the previous results, suggesting that prophylactic administration of antibiotics did not interfere with procedure-related infection rate (Table 4). Three studies<sup>29,35,38</sup> reported the anticoagulant profiles, and results showed that thromboprophylaxis did not influence the risk for thrombosis either. In the subgroup analysis of insertion methods, only one study<sup>31</sup> used surgical cut-down approach in all patients, and six studies<sup>10,33–36,40</sup>

Table 2 Risk of bias of included randomized controlled trials based on Cochrane Handbook for Systematic Reviews of Interventions

Study	Sequence generation	Allocation concealment	Blinding	Incomplete outcome data	Selective out- come reporting	Other sources of bias	Risk of bias
D'Angelo et al, 2002 <sup>30</sup>	Yes	Uncertain	Uncertain	Yes	Yes	Yes	Moderate
Biffi et al, 2009 <sup>29</sup>	Yes	Yes	Uncertain	Yes	Yes	Yes	Low

Notes: Low risk: 5–6 sections with "yes"; moderate risk: 3–4 sections with "yes"; high risk: ≤2 sections with "yes".

Study	Representativeness of the exposed cohort	Selection of the non- exposed	Ascertainment of exposure	Demonstration that outcome of interest was not present at start of study	Comparability of cohorts on the basis of the design or	Assessment of outcome	Was follow- up long enough for outcomes to	Adequacy of follow- up of cohorts	Quality score
							0000		
Alahyane et al, 2010 <sup>32</sup>	_	_	0	_	2	_	0	0	6
Akahane et al, 2011 <sup>33</sup>	_	_	_	_	2	_	_	_	6
Goltz et al, 2012 <sup>34</sup>	_	_	_	_	2	_	0	0	7
Goltz et al, 2013 <sup>10</sup>	_	_	_	_	2	_	_	_	6
Kuriakose et al, 2002 <sup>35</sup>	_	_	_	_	2	_	_	0	80
Li et al, 2016 <sup>36</sup>	_	_	_	_	2	_	_	_	6
Marcy et al, 2005 <sup>37</sup>	_	_	_	_	2	_	_	_	6
Marcy et al, 2008 <sup>38</sup>	_	_	_	_	2	_	_	0	80
Matiotti-Neto et al, 2017 <sup>39</sup>	_	_	_	_	_	_	0	0	6
lorio et al, 2018 <sup>31</sup>	_	_	_	_	2	_	0	0	7
Shiono et al, 2014 <sup>40</sup>	_	_	_	_	2	_	0	0	7
Notes: Otality of observational study is sessed according to Nowrestia-Otrawa scale (NOS) consisting of three commants: nation (4 noints) commanalility of the study sesses according to Norses and sessesment of nirrow a Ranine)	tudv is assessed according to N	ewcastle-Ottawa	scale (NOS) consisting o	of three components, patient se	laction (4 points) compar	ahility of the study an	ours (2 nointe) and as	seesement of outo	me (3 nointe)

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employed puncture approach for all patients. The results did not change the conclusions for the primary outcomes from original analyses, and pooled data from six studies including 1,781 patients who underwent percutaneous puncture approach indicated consistent results showing that the incidences of intra-operative events in chest port group were significantly higher (2.05% and 0.12%, OR 7.87, 95% CI 1.82–34.13; p<0.01) (Table 4). Finally, two subgroups were stratified according to patients' age group (<60 years old and  $\geq$ 60 years old). After pooling data from eight studies including 2,063 patients less than 60 years old, the result showed that no significant difference was detected with regard to intra-operative events between chest ports and arm ports (1.19% and 0.02%, OR 2.59, 95% CI 0.74–9.08; p<0.14), suggesting that age may be a risk factor for intra-operative events (Table 4).

#### Sensitivity analysis and publication bias

One low risk RCT<sup>29</sup> and six non-randomized comparative studies<sup>10,33,35–38</sup> that scored over eight points according to NOS were included in the sensitivity analysis. The results of sensitivity analysis did not change the conclusions for total complication events, all of the primary outcomes, and incidence of procedure conversion from original analyses. However, between-study heterogeneity decreased slightly in all of these parameters (Table 5). Due to insufficient data, sensitivity analyses for early port removal and operating time were not performed.

Figure 5 depicts a funnel plot of included studies that assessed primary outcomes of interest. The distribution was even and around the vertical. Most of the studies were within the 95% CIs, suggesting no apparent publication bias.

#### Discussion

This systematic review with meta-analysis compared the clinical efficacy of two types of TIVAPs: chest port and arm port. Our results showed that arm port is a safe site for implantation, with lower incidence of intra-operative complications, however, arm port is associated with higher procedure conversion rate. No statistically significant differences between chest port and arm port were found in total complication events, procedure-related infections, procedure-related thrombotic complications, incidence of early port removal, and operating duration. On subgroup analysis, we found that antibiotic prophylaxis and method of insertion did not interfere with the conclusions for the primary outcomes from original analyses, but results revealed that there were no statistically significant



Figure 2 Forest plot and meta-analysis of total complication events. Abbreviation: M-H, Mantel-Haenszel method.

differences in intra-operative events in those patients under 60, suggesting that age may affect the risk for intraoperative complications. Sensitivity analysis did not change the conclusions of all endpoints of interest.

For the utilization of a new technique, patients' safety is always the first priority. Chest ports are still popular and their application is continuously expanding among nonsurgical physicians.<sup>15</sup> Under the guidance of ultrasound or fluoroscopy, the risks for intra-operative events, such as pneumothorax, hemothorax, and arterial injury, are therefore minimized during percutaneous cannulations of subclavian vein or jugular vein.<sup>42</sup> Implantation of arm ports through peripheral forearm vessels is more surgically challenging, but arm port users experience less perception of "foreign body", with satisfactory cosmetic results. Pooled data from 2,176 chest port and 1,720 arm port users from included studies of this study, showed that incidences for total complication events were 11.8% and 12.8%, respectively. Our result of total complication events is in agreement with previous studies that assessed overall complication rate, 4,19,43 with no significant difference between chest ports and arm ports. Although the technique of implanting arm ports is newly developed, and more challenging for clinicians, our findings indicate that arm ports are at least as safe as conventional chest ports with proper patient indications.

The procedure conversion rate is significantly higher for arm ports as compared to chest ports. Additional challenges for arm port implantation include anatomic variations, longer distance for implantation, and frequent use of peripheral intravenous access, which make cannulation of peripheral forearm vessels not always achievable. Anatomic variations of basilic, cephalic or brachial vein frequently occur, and these peripheral vessels are much smaller in diameter as compared with central veins. These peripheral forearm vessels may be difficult to recognize or too limited in size to use.<sup>44</sup> Therapeutic usage of peripheral venous access is frequent for oncologic and critically-ill patients, however, frequent puncture of these vessels leads to adhesion, spasm, or shrinkage, and thus complicates anatomy of surrounding structures. Moreover, patients with conversion might reveal higher risks for other complications. As implantation of indwelling TIVAPs is performed not only by surgeons, but also widely performed by non-surgical practitioners (such as trained physicians, radiologists, anesthesiologists, oncologists, and so forth), arm port implantation may need to be placed in selected patients to avoid procedure conversion. In our experience, no matter how experienced the clinician is in arm port implantation, preoperative ultrasound evaluation of forearm vessels is indicated in every case to minimize this unfavorable event.

Our findings showed that chest port was associated with higher incidence of intra-operative complications. Due to specific anatomy, accidental pneumothorax or arterial puncture could not be eliminated completely, even if ultrasound or fluoroscopy guidance is routinely available. But interestingly, in subgroup analysis, no between-group differences in intra-operative events were seen in those patients under 60. Patients' age appeared to be relevant to the risk for intra-operative events, although it is still poorly illustrated. Similar results were previously reported: high age is an independent risk factor for procedural complications in subclavian vein puncture, and one implication is closely related to variations in body

Study or subgroup		ort	arm p			Odds ratio	Odds ratio
		Total	Events	Total	Weight	M-H, Fixed, 95% Cl	M-H, Fixed, 95% Cl
1.2.1 procedure related inf	ections						
Akio-akhane 2011	4	47	9	115	10.8%	1.10 [0.32, 3.75]	
Biffi 2009	4	270	3	133	9.0%	0.65 [0.14, 2.95]	
Goltz 2012	3	52	9	152	9.8%	0.97 [0.25, 3.74]	
Goltz 2013	1	25	1	25	2.2%	1.00 [0.06, 16.93]	
Kuriakose 2002	8	273	. 9	149	25.6%	0.47 [0.18, 1.24]	
Li 2016	8	237	1	107	3.0%		
						3.70 [0.46, 29.99]	
Marcy 2005	5	100	2	100	4.3%	2.58 [0.49, 13.62]	
Marcy 2008	3	112	4	113	8.8%	0.75 [0.16, 3.43]	
Mariomatiottio neto 2017	6	247	1	195	2.5%	4.83 [0.58, 40.46]	
Olga lorio 2018	1	106	2	109	4.4%	0.51 [0.05, 5.70]	
Shiono 2014	14	342	8	257	19.8%	1.33 [0.55, 3.22]	
Subtotal (95% CI)		1811	-	1455	100.0%	1.11 [0.74, 1.66]	•
Total events	57		49				Ē
		0.00.1					
Heterogeneity: <i>Chi</i> ²=8.42, <i>d</i> Test for overall effect: <i>Z</i> =0.4	•		-=0%				
1.2.2 procedure related thr	ombotic	compli	cations				
Akio-akhane 2011	0	47	0	115		Not estimable	
Biffi 2009	23	270	11	133	26.5%	1.03 [0.49, 2.19]	-+
Goltz 2012	2	52	21	152	20.3%	0.25 [0.06, 1.10]	
Goltz 2012	1	25	1	25	1.9%	1.00 [0.06, 16.93]	
Kuriakose 2002	5	273	3	149	7.5%	0.91 [0.21, 3.85]	
Li 2016	9	237	1	107	2.6%	4.18 [0.52, 33.45]	
Marcy 2005	3	100	1	100	1.9%	3.06 [0.31, 29.95]	
Marcy 2008	3	112	12	113	22.9%	0.23 [0.06, 0.84]	
Mario matiottio neto 2017	1	247	4	195	8.8%	0.19 [0.02, 1.75]	
Olga lorio 2018	1	106	0	109	1.0%	3.11 [0.13, 77.29]	
Shiono 2014	4	342	3	257	6.7%	1.00 [0.22, 4.52]	
Subtotal (95% CI)	-	1811			100.0%	0.75 [0.49, 1.13]	•
	60	1011	67	1455	100.070	0.75 [0.45, 1.15]	•
Total events Heterogeneity: <i>Chi</i> ²=12.52,	52		57				
1.2.3 intra-operative events	s			145	2.4.00	40.00 /0.00 000 501	
Alde aldeana 2044	2						
Akio-akhane 2011	2	47	0	115		12.69 [0.60, 269.53]	
Biffi 2009	0	270	1	133	22.2%	0.16 [0.01, 4.04]	·
		270 52		133 152			· · · · · · · · · · · · · · · · · · ·
Biffi 2009	0	270	1	133		0.16 [0.01, 4.04]	· · · · · · · · · · · · · · · · · · ·
Biffi 2009 Goltz 2012	0 0	270 52	1 0	133 152		0.16 [0.01, 4.04] Not estimable	· · · · · · · · · · · · · · · · · · ·
Biffi 2009 Goltz 2012 Goltz 2013	0 0 0	270 52 25	1 0 0	133 152 25		0.16 [0.01, 4.04] Not estimable Not estimable Not estimable	<
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016	0 0 0	270 52 25 273	1 0 0 0	133 152 25 149	22.2%	0.16 [0.01, 4.04] Not estimable Not estimable Not estimable 6.15 [0.79, 47.64]	· · · · · · · · · · · · · · · · · · ·
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005	0 0 0 13 0	270 52 25 273 237 100	1 0 0 1 0	133 152 25 149 107 100	22.2%	0.16 (0.01, 4.04) Not estimable Not estimable Not estimable 6.15 (0.79, 47.64) Not estimable	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008	0 0 0 13 0 1	270 52 25 273 237 100 112	1 0 0 1 0	133 152 25 149 107 100 113	22.2% 14.4% 5.4%	0.16 [0.01, 4.04] Not estimable Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77]	
Biffi 2009 Goltz 2012 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017	0 0 13 1 4	270 52 273 237 100 112 247	1 0 0 1 0 4	133 152 25 149 107 100 113 195	22.2%	0.16 [0.01, 4.04] Not estimable Not estimable 8.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018	0 0 13 0 1 4 0	270 52 25 273 237 100 112 247 106	1 0 0 1 0 4 0	133 152 25 149 107 100 113 195 109	22.2% 14.4% 5.4% 48.7%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014	0 0 13 1 4	270 52 25 273 237 100 112 247 106 342	1 0 0 1 0 4	133 152 25 149 107 100 113 195 109 257	22.2% 14.4% 5.4% 48.7% 6.2%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI)	0 0 13 0 1 4 0 5	270 52 25 273 237 100 112 247 106	1 0 0 1 0 4 0 0	133 152 25 149 107 100 113 195 109 257	22.2% 14.4% 5.4% 48.7%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga lorio 2018 Shiono 2014 Subtotal (95% CI) Total events	0 0 13 0 1 4 0 5 25	270 52 25 273 237 100 112 247 106 342 <b>1811</b>	1 0 1 0 4 0 0 6	133 152 25 149 107 100 113 195 109 257	22.2% 14.4% 5.4% 48.7% 6.2%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi<sup>z</sup>=</i> 7.82, <i>d</i>	0 0 0 13 0 1 4 5 5 7=5 (P=0.	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*=	1 0 1 0 4 0 0 6	133 152 25 149 107 100 113 195 109 257	22.2% 14.4% 5.4% 48.7% 6.2%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga lorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1	0 0 0 13 13 1 4 0 5 f=5 (P=0. 3 (P=0.0) 3 (P=0.0)	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3)	1 0 0 1 0 4 0 0 0 8 36%	133 152 25 149 107 100 113 195 109 257 1455	22.2% 14.4% 5.4% 48.7% 6.2% 100.0%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi<sup>=</sup>=</i> 7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio- akhane 2011	0 0 0 13 1 4 0 5 25 7=5 (P=0. 3 (P=0.0) 3 (P=0.0) 0	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3)	1 0 0 1 0 4 0 0 36% 6	133 152 25 149 107 100 113 195 109 257 1455	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio-akhane 2011 Biffi 2009	0 0 0 13 0 1 4 0 5 5 5 5 (P=0. 3 (P=0.0) 3 (P=0.0) 10	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3) 47 270	1 0 0 1 0 4 0 0 8 36% 8 8	133 152 25 149 107 100 113 195 109 257 1455 115 133	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi<sup>=</sup>=</i> 7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio- akhane 2011	0 0 0 13 1 4 0 5 25 7=5 (P=0. 3 (P=0.0) 3 (P=0.0) 0	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3)	1 0 0 1 0 4 0 0 36% 6	133 152 25 149 107 100 113 195 109 257 1455	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio-akhane 2011 Biffi 2009	0 0 0 13 0 1 4 0 5 5 5 5 (P=0. 3 (P=0.0) 3 (P=0.0) 10	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3) 47 270	1 0 0 1 0 4 0 0 8 36% 8 8	133 152 25 149 107 100 113 195 109 257 1455 115 133	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario 2008 Mario matiottio neto 2017 Olga lorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi<sup>2</sup></i> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio-akhane 2011 Biffi 2009 Goltz 2012	0 0 0 13 0 1 4 0 5 5 5 ( <i>P</i> =0.0) 3 ( <i>P</i> =0.0) 3 ( <i>P</i> =0.0) 10 1	270 52 25 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 3) 47 270 52	1 0 0 1 0 4 0 0 36% 6 8 0	133 152 25 149 107 100 113 195 109 257 1455 115 133 152	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 5.43 [0.25, 118.96]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 <b>Subtotal (95% CI)</b> Total events Heterogeneity: <i>Chi<sup>=</sup>=</i> 7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio- akhane 2011 Biffi 2009 Goltz 2013 Kuriakose 2002	0 0 0 13 0 1 4 0 5 5 (P=0.0 3 (P=0.0) 3 (P=0.0) 10 10 12 27	270 52 255 273 237 100 112 247 106 342 1811 17); /*= 3) 47 270 52 25 273	1 0 0 0 4 0 0 6 36% 6 8 0 0 14	133 152 25 149 107 100 113 195 109 257 1455	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.9% 33.9%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 5.43 [0.25, 118.96] 1.06 [0.54, 2.09]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario matiottio neto 2017 Olga Iorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio-akhane 2011 Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016	0 0 0 13 0 1 4 0 5 7=5 (P=0. 3 (P=0.0: 3 (P=0.0: 10 10 1 2 27 25	270 52 255 273 237 100 112 247 106 342 1811 17); /*= 3) 47 270 52 52 273 237	1 0 0 1 0 4 0 0 6 36% 6 8 0 14 7	133 152 25 149 107 100 113 195 109 257 1455 133 152 25 149 107	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5% 0.5% 0.9% 33.9% 17.9%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 5.43 [0.25, 118.96] 1.06 [0.54, 2.09] 1.68 [0.70, 4.03]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario 2008 Mario matiottio neto 2017 Olga lorio 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi<sup>2</sup>=7.82, d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Aktio-akhane 2011 Biffi 2009 Goltz 2012 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005	0 0 0 13 0 1 4 0 5 7=5 (P=0.0 3 (P=0.0) titions 0 10 1 2 27 25 5	270 52 273 237 100 112 247 106 112 247 1811 17); /²= 1811 17); /²= 270 52 253 237 100	1 0 0 4 0 36% 6 36% 6 8 0 0 14 7 5	133 152 25 149 107 100 113 195 257 1455 133 152 255 149 257 107 100	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5% 0.9% 33.9% 17.9%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 <b>[1.07, 5.29]</b> 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 5.43 [0.25, 118.96] 1.66 [0.54, 2.09] 1.68 [0.74, 4.03] 1.00 [0.28, 3.57]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario 2008 Shiono 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio- akhane 2011 Biffi 2009 Goltz 2012 Goltz 2012 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005	0 0 0 13 0 1 4 0 5 5 7=5 (P=0. 3 (P=0.0) tions 0 10 1 2 27 25 5 3	270 52 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 187 270 52 273 237 270 52 275 273 237 100 112	1 0 0 4 0 0 4 0 0 6 6 8 8 0 1 4 7 5 2	133 152 25 149 107 100 113 195 109 257 1455 133 152 25 149 107 100 113	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5% 0.5% 0.9% 33.9% 17.9% 9.9% 4.0%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 1.06 [0.54, 2.09] 1.68 [0.70, 4.03] 1.00 [0.28, 3.57] 1.53 [0.25, 9.32]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017 Olga lorio 2018 Shiono 2014 <b>Subtotal (95% CI)</b> Total events Heterogeneity: <i>Chi<sup>=</sup>=7.82, d</i> Test for overall effect: <i>Z=2.1</i> <b>1.2.4 mechanical complica</b> Akio-akhane 2011 Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Marcy 2008 Mario matiottio neto 2017	0 0 0 13 0 1 4 0 5 5 7=5 (P=0. 3 (P=0.0) 3 (P=0.0) 10 10 10 27 25 5 3 3 3	270 52 255 273 237 100 112 247 106 342 1811 17); /= 2 3) 47 270 52 25 273 237 100 52 25 273 237 100 52 25 273 237	1 0 0 1 0 0 0 0 0 0 0 0 0 0 0 0 0 14 7 5 2 0 0	133 152 25 149 107 100 113 195 1455 1455 1455 1455 149 107 100 113 195	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5% 0.9% 33.9% 17.9%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 22.1.48] 5.43 [0.25, 118.96] 1.06 [0.54, 2.09] 1.68 [0.70, 4.03] 1.00 [0.28, 3.57] 1.53 [0.25, 9.32] 5.60 [0.29, 109.01]	
Biffi 2009 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005 Mario 2008 Shiono 2018 Shiono 2014 Subtotal (95% CI) Total events Heterogeneity: <i>Chi</i> <sup>2</sup> =7.82, <i>d</i> Test for overall effect: <i>Z</i> =2.1 <b>1.2.4 mechanical complica</b> Akio- akhane 2011 Biffi 2009 Goltz 2012 Goltz 2012 Goltz 2012 Goltz 2013 Kuriakose 2002 Li 2016 Marcy 2005	0 0 0 13 0 1 4 0 5 5 7=5 (P=0. 3 (P=0.0) tions 0 10 1 2 27 25 5 3	270 52 273 237 100 112 247 106 342 <b>1811</b> 17); /*= 187 270 52 273 237 270 52 275 273 237 100 112	1 0 0 4 0 0 4 0 0 6 6 8 8 0 1 4 7 5 2	133 152 25 149 107 100 113 195 109 257 1455 133 152 25 149 107 100 113	22.2% 14.4% 5.4% 48.7% 6.2% 100.0% 7.8% 21.5% 0.5% 0.5% 0.9% 33.9% 17.9% 9.9% 4.0%	0.16 [0.01, 4.04] Not estimable Not estimable 6.15 [0.79, 47.64] Not estimable 3.05 [0.12, 75.77] 0.79 [0.19, 3.18] Not estimable 8.39 [0.46, 152.47] 2.38 [1.07, 5.29] 0.18 [0.01, 3.21] 0.60 [0.23, 1.56] 8.88 [0.36, 221.48] 1.06 [0.54, 2.09] 1.68 [0.70, 4.03] 1.00 [0.28, 3.57] 1.53 [0.25, 9.32]	
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#### Figure 3 Forest plot and meta-analysis of primary outcomes. Abbreviation: M-H, Mantel-Haenszel method.

habitus.45,46 Based on our experiences, our explanation is that vessel fragility and decreased vessel elasticity may be responsible for this issue, as the elderly are prone to compromised vascular compliance, which makes the

aged population more susceptible to vascular interference and trauma. Another potential reason is higher morbidities of chronic obstructive pulmonary diseases and obesity that attribute to structural elevation of apex pulmonis, thereby



Figure 4 Forest plot and meta-analysis of secondary outcomes. (A) Comparison of procedure conversion rate and early port removal. (B) Comparison of operating time between groups.

Abbreviations: M-H, Mantel-Haenszel method; IV, inverse variance method.

facilitating the occurrence of pneumothorax.<sup>46</sup> Certainly, special caution in terms of intra-operative complications in aged patients is alerted when performing implantation of chest ports.

The pooled analyses of procedure-related infections, thrombolytic complications, and mechanical complications showed no between-group differences. Subgroup analysis of antibiotic prophylaxis demonstrated that administration of antibiotics did not affect the outcome of overall infection rate. But we still recommend that anti-septic protocols should be followed for the sake of infection prevention and management in oncologic patients who are vulnerable to infection owing to leuimmune suppression.47 kocytopenia induced by Subgroup analysis of thromboprophylaxis did not alter any original conclusion, which was consistent with Chaukiyal's result<sup>48</sup> showing that thromboprophylaxis had no beneficial effects on the risks of catheterrelated thrombolytic complications in oncologic patients. Nonetheless, as regular heparin flushing has become routine practice for TIVAPs in almost every institution,<sup>49</sup> it seems to be enough for the prevention of port-related thrombosis.50

Two methods of implanting TIVAPs are used in current clinical practice: surgical cut-down or percutaneous puncture. No consensus has ever been reached regarding implantation methods because percutaneous puncture is less timeconsuming, but surgical cut-down method lowers the risk for complications such as pneumothorax.<sup>31</sup> With the worldwide spread of ultrasound monitoring for venous access. venous surgical cut-down method is on the decrease.<sup>31</sup> No significant difference in terms of insertion approach could be addressed in this study. Of the three included studies in which operating time was reported, detailed procedure description was not available in one study,<sup>30</sup> one study<sup>31</sup> inserted TIVAPs through surgical cut-down, and the last<sup>39</sup> separately used surgical approach and puncture to cannulate arm ports and chest ports, respectively. Because of the heterogeneous characteristic and data insufficiency, it is difficult to reach a definite conclusion for the comparison of inserting methods in this instance.

Arm port implantation is especially attractive to young females for cosmetic reasons, as it leaves no scars on the chest. Arm port users experience less "bra inconvenience", and psychologically, patients' anxiety of indwelling arm ports might be less as compared to chest ports.<sup>40</sup>

Group	Pro	Procedure-related infections	ed infe	ections	Proc	Procedure-related thrombotic complications	d throi	mbotic	Intr	Intra-operative events	ents		Mec	Mechanical complications	licatio	us
	2	OR(95% CI)	ا <sup>2</sup> (%)	Heterogeneity	۲	OR(95% CI)	1 <sup>2</sup> (%)	Heterogeneity	2	OR(95% CI)	1 <sup>2</sup> (%)	Heterogeneity	2	OR(95% CI)	ا <sup>2</sup> (%)	Heterogeneity
Overall	=	1.11 (0.74–1.66)	0	0.59	=	0.75 (0.49–1.13)	28	0.19	=	2.38 (1.07–5.29)	36	0.17	=	1.34 (0.92–1.95)	25	0.21
Antibiotic prophylaxis																
Yes	m	1.12	0	0.74	N/A				N/A				A/N			
No	3	(0.56–2.24) 1.24 (0.55–2.76)	0	0.52	N/A				N/A				N/A	_		
Anti-coagulant prophylaxis																
Yes	N/A				_	0.91	Ż	N/A	N/A				N/A			
٥ Z	N/A	~			2	(0.21–3.85) 0.54 (0.12–2.32)	A 74	0.05	N/A				N/A	·		
Method of insertion																
Surgical cut-down	_	0.51	Ż	N/A	-	3.11	Ż	N/A	-	0	Ż	N/A	-	0	Ż	N/A
Puncture	6	(0.05–5.70) 1.03 (0.63–1.68)	< 0	0.51	6	(0.13–77.29) 0.81 (0.42–1.56)	A 19	0.29	6	7.87 (1.82–34.13)	۷ ٥	0.93	6	1.55 (0.98–2.47)	38 38	0.15
<b>Age</b> <60 years old	œ	0.90	0	0.6	ω	0.78	35	0.15	œ	2.59	4	0.17	œ	4	0	0.51
>60 years old	2	(0.54–1.52) 1.25 (0.61–2.54)	0	0.8	2	(0.50–1.23) 1.00 (0.22–4.52)	Ż∢	N/A	2	(0.74–9.08) 9.81 (1.13–85.37)	0	0.84	2	(0.75–1.74) 1.47 (0.03–71.26)	79	0.03
Abbreviations: n, number of studies; N/A, not applicable.	ber of	studies; N/A, no	ot applic	able.	]				]		1					

Outcome of interest	Number	Number of chest	Number of arm	OR (95%CI)	p-value	1 <sup>2</sup>	Heterogeneity
	of studies	port patients	port patients			(%)	
Total complication events	7	1,064	742	0.88 (0.67–1.17)	0.38	41	0.12
Procedure-related	7	1,064	742	0.95 (0.56–1.62)	0.86	0	0.49
infections							
Procedure-related	7	1,064	742	0.92 (0.56–1.51)	0.74	34	0.18
thrombotic complications							
Intra-operative events	7	1,064	742	3.28 (1.03-10.40)	0.04	33	0.21
Mechanical complications	7	1,064	742	1.06 (0.71–1.58)	0.79	0	0.52
Conversion rate	3	619	353	0.33 (0.19–0.57)	<0.001	0	0.44

Table 5 Sensitivity analysis comparing chest ports and arm ports



Figure 5 Funnel plot demonstrating meta-analysis of primary outcomes of interest. Abbreviation: SE, standard error.

Moreover, arm port is especially appropriate for patients who require neck or chest radiation therapies.<sup>40</sup> Comparison of these quality-of-life related outcomes is of importance, however, only a limited number of studies reported these parameters, which were measured based on various evaluating methods, such as self-administered questionnaire<sup>10,36</sup> and quality of life score,<sup>37,46</sup> and non-uniformity of these measurements made these data incomparable in this study.

Our study is distinctive and provides valuable information to oncology-related clinicians in that, to the best of our knowledge, this systematic review with meta-analysis is the first one to comprehensively investigate the clinical efficacies and represents the most up-to-date information with respect to chest ports and arm ports. We used multiple strategies to launch evidence acquisition, strict standards to assess quality of included studies, scientific statistics for data analysis, subgroup and sensitivity analysis to control study heterogeneity. Our study has several limitations which should be taken into consideration. First of all, the most apparent limitation of the present study is that most of the included data were from retrospective cohort studies. Inadequacy of randomization and lack of blinding will increase the risk of bias. Secondly, this study was based on the premised assumption that demographic subgroups (age, sex, race, nationality, and disease distribution) were similar enough for comparison. However, subgroup analysis with regard to age generated some different outcomes, hence, further systematic reviews should be conducted when sufficient future results are released. And then, the included studies were from different institutions with various levels of clinician expertise. Not only the between-study time-span, which is more than 10 years, but also the different backgrounds of the TIVAP operators with different insertion approaches will affect the outcomes of interest. Next, some outcomes of interest, especially the mechanical complications, might not be clearly defined in some studies, yielding misinterpretation of results or data omission. So, the incidence of mechanical complications might be underrated, and caution should be taken when interpreting results of this section. Finally, follow-up period of included studies was generally short. But there are increasing cumulative risks for TIVAP-related infection, thrombosis, and mechanical mis-functioning along with the duration of port indwelling, some incidences of outcomes might be underestimated because of insufficient follow-up period. Surely, long-term outcomes of arm ports, especially for oncologic efficacy and safety, require further investigation.

#### Conclusion

In this systematic review with meta-analysis that compared clinical efficacy of two types of TIVAPs: chest port and arm port, we found that arm port is associated with higher procedure conversion rate, but lower incidence of intraoperative complications. There are no statistically significant differences between chest port and arm port with regard to in total complication events, procedure-related infections, procedure-related thrombotic complications, incidence of early port removal, and operating duration. Age may be a risk factor for intra-operative complications. Further confirmation of these results is warranted and our findings should be updated with larger scale and well-designed studies with longer follow-up.

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# Disclosure

The authors report no conflicts of interest in this work.

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