

Laparoscopic versus open sacrocolpopexy for treatment of prolapse of the apical segment of the vagina: a systematic review and meta-analysis

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Abstract

Introduction Pelvic organ prolapse is showing an increasing prevalence (3 – 50 %). The gold standard treatment of apical prolapse is sacrocolpopexy which can be performed via minimal access (laparoscopy or robotics) or open approaches. The aim of this review was to appraise the effectiveness of minimal access surgery versus the open approach in the treatment of apical prolapse.

Methods Keywords were searched in: CINAHL, MEDLINE, CENTRAL, Cochrane MDSG Trials Register, Cochrane Library, Current Controlled Trials, ClinicalTrials.gov, WHO International Trials Registry Platform search portal, LILACS, and Google Scholar databases. Data up to 31 April 2014 were considered. Randomized and nonrandomized controlled trials evaluating all women who underwent minimally invasive sacropexy (MISC) and open sacropexy (OSC) were included. A data extraction tool was used for data collection. MISC was compared with OSC using narrative analysis and meta-analysis (RevMan) where appropriate.

Results MISC and OSC were compared in 12 studies involving 4,757 participants. MISC and OSC were equally effective in terms of point-C POP-Q measurements and recurrence rate.

MISC was associated with a lower transfusion rate (odds ratio 0.41, 95 % CI 0.20 – 0.83), shorter length of hospital stay (mean difference –1.57 days, 95 % CI –1.91 – –1.23 days), and less blood loss (mean difference –113.27 mL, 95 % CI –163.67 – –62.87 mL) but a longer operating time (mean difference 87.47, 95 % CI 58.60 – 116.34, $p < 0.0001$).

Conclusions MISC showed similar anatomic results to OSC with a lower transfusion rate, shorter length of hospital stay and less blood loss. The rate of other complications was similar between the approaches. Cautious interpretation of results is advised due to risk of bias caused by the inclusion of nonrandomized studies.

Keywords Apical prolapse · Laparoscopy · Robotic surgery · Sacrocolpopexy

Introduction

Pelvic organ prolapse has a prevalence of 3 – 6 % when based on symptoms and up to 50 % when based on vaginal examination [1]. Specifically, prevalence of apical prolapse ranges between 0.2 % and 43 % [2]. In the UK it represents approximately 1.1 % of the NHS (National Health Service) budget [3]. In England, 16.9 % of women were admitted in 2009 for treatment of pelvic organ prolapse, representing a cost of €81,030,907 [4].

Open sacrocolpopexy is considered the gold standard surgical treatment of apical prolapse [5]. However, it is associated with longer time to return to daily activities, longer operating times (OT) and greater costs [5]. Therefore, laparoscopic surgery has gained support amongst surgeons because of its advantages, but its marked learning curve has limited its adoption. Conversely, robotic surgery has arisen as the latest technology capable of offering benefits in terms of dexterity and shortening the learning curve [6].

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The aim of this review was to appraise the effectiveness and safety of minimal access surgery in relation to the open approach in the treatment of apical prolapse to determine the best approach to performing sacropexy. Currently, there are three systematic reviews covering a similar topic. The first [7] included observational studies up to 2010, and the second was a Cochrane review [8]. From this date onwards, a variety of new research studies have been completed, which justifies an update on this topic. Additionally, just one and two studies, respectively, related to sacropexy were included in each review, and this may raise doubts concerning transferability [9]. Moreover, neither of these reviews compared minimally invasive sacropexy (MISC) and open sacropexy (OSC). The third systematic review [10] focused on robotic sacropexy (RSC) and did not include (among other studies) a recently published randomized controlled trial (RCT) comparing RSC and laparoscopic sacropexy (LSC). Additionally, this third review also included uncontrolled studies which are more susceptible to bias [11].

Objectives of the review

- To compare the effectiveness of MISC and OSC in the treatment of apical prolapse.
- To appraise the outcomes of MISC and OSC in relation to intraoperative and postoperative complications, mortality, postoperative length of stay (LOS), postoperative pain, estimated blood loss (EBL), OT and quality of life (QoL).

Research question

Which approach (MISC or OSC) is more effective when performing a sacropexy to treat women with prolapse of the apical segment of the vagina?

Methods

Selected databases were systematically searched using relevant key words (Table 1) to identify pertinent studies. The selected databases included: Cumulative Index to Nursing and Allied Health Literature (CINAHL), MEDLINE, Cochrane Central Register of Controlled Trials

(CENTRAL), Cochrane Menstrual Disorders and Subfertility Group (MDSG) Trials Register, Cochrane Library, Current Controlled Trials, ClinicalTrials.gov, WHO International Trials Registry Platform search portal, Latin American and Caribbean Health Science Literature (LILACS), and Google Scholar. Additionally, the reference list of relevant studies, the *International Urogynecology Journal* and the *Journal of Minimally Invasive Gynecology* [9] were also searched by hand.

Duplicate titles and abstracts were removed and the remaining studies were selected according to relevance and the use of a flow diagram. Abstracts were assessed against the inclusion and exclusion criteria and reasons for exclusion were documented. Two authors assessed selected full-text papers and those that did not meet the inclusion criteria were excluded. Finally, chosen papers were appraised rigorously to determine their quality using appropriate tools.

Criteria for study selection

Inclusion criteria

- All women with prolapse of the apical segment of the vagina who underwent OSC or MISC (RSC or LSC)
- Studies written in English, Portuguese, or Spanish
- Studies from 1980 to 2014
- Quantitative studies including experimental studies (RCTs), quasiexperimental studies (controlled before-and-after studies), and controlled observational studies (cohort studies)
- Studies evaluating effectiveness of sacropexy, complications, LOS, EBL, OT, mortality, postoperative pain or postoperative QoL

Exclusion criteria

- All women who underwent a different procedure for the surgical repair of apical prolapse
- Studies not written in English, Portuguese, or Spanish
- Case control studies, uncontrolled before-and-after studies, cross-sectional surveys, case series
- Studies in cadavers

Table 1 Relevant key words

Key concepts	Alternative terms	Boolean Operators
Sacrocolpopexy	Sacral colpopexy, sacral promontofixation, colposacral suspension, sacropexy, colpopexy	AND
Prolapse of the apical segment of the vagina	Vaginal vault prolapse, apical prolapse	OR
Robotic	Robot, robot*, robot-assisted, minimal access surgery, minimally invasive surgery	AND
Laparoscopy	Laparoscop*, minimal access surgery, minimally invasive surgery, keyhole surgery	OR
Abdominal sacrocolpopexy	Open sacrocolpopexy	AND

Quality assessment

The criteria to determine whether the included studies had low, high or unclear risk of bias were based on the appraisal tool recommended by the Cochrane Collaboration [9], which explores selection bias, performance bias, detection bias, attrition bias and reporting bias. Other biases were also considered: whether funding and conflicts of interests were reported in the included studies [9]. The Newcastle-Ottawa scale (NOS) was also used for quality assessment of nonrandomized studies because it provides an objective approach to the evaluation of validity that can be fully reported [9]. It evaluates eight items and each one can be awarded one star, except the item “comparability” that can be awarded two stars (maximum of nine). The NOS was customized to this research choosing two relevant confounders. “Previous abdominal surgery” was selected because it is an independent predictor of potential complications after surgery [12] whereas “concomitant surgery” is a variable related to longer OT [13]. Also, it was decided that the period of follow-up should be no less than 12 months to allow time to detect prolapse recurrence [14]. The maximum acceptable percentage of subjects lost to follow-up was agreed to be 20 % [15].

Data extraction

A data extraction tool was developed based on the aim, objectives and research question of this review. Effectiveness was defined as point-C POP-Q measurement at more than 1 cm above the level of the hymen [16]. Subjective effectiveness was defined as the woman reporting no symptoms after the procedure [16]. Other outcomes of interest were: LOS, EBL, OT and postoperative QoL. Complications and mortality were also recorded to assess the safety of the procedure. Authors were contacted by email to acquire missing information but no responses were obtained.

Data analysis

Narrative analysis and meta-analysis were used. For studies with different designs, narrative analysis was implemented using tables [17]. Review Manager 5.3 was used for meta-analysis [17]. For categorical outcomes, the odds ratio (OR) was calculated using the Mantel-Haenszel method [9]. For continuous variables, the mean difference (MD) was derived from means and standard deviations and used when outcomes were reported using identical scales. When scales were different, the standardized MD (SMD) was derived using RevMan [9]. The confidence interval was set at 95 % and p values <0.05 (two-tailed) were considered statistically significant. Papers in which authors reported outcomes using medians and ranges were not included in the meta-analysis [9].

RevMan was used to calculate heterogeneity using the chi-squared test and the percentage variability (I^2). When heterogeneity was lower than 50 %, fixed-effects meta-analysis was performed. When heterogeneity was equal to or higher than 50 %, heterogeneity was explored (i.e. for possible causes of heterogeneity) and, additionally, a random-effects meta-analysis was undertaken [9]. Subgroup analyses were performed where appropriate. A sensitivity analysis was also performed. This involved inclusion of high-quality studies (i.e. including RCTs and excluding observational studies NOS scores of five or less), reconsideration of the methods used for analysis (appropriate usage of fixed-effects versus random-effects methods, or MD versus SMD) and making assumptions on how missing data could have affected the results (assigning lost subjects to the worst case scenario) [9, 11].

Results

Details of studies

Full reports of 28 studies were appraised. The process of screening and selection is illustrated using a PRISMA flow chart (Fig. 1). After assessment of 28 studies, eight were excluded and the reasons are summarized in Table 2.

Two comparisons were made: MISC (i.e. RSC plus LSC) versus OSC which is presented in this review, and RSC versus LSC which is discussed in another review. Papers were therefore divided accordingly. From 20 studies included in the analysis, 12 compared MISC and OSC (Table 3) and eight RSC and LSC. However, the study by Nosti et al. [12] also compared RSC and LSC, and thus was included in both analyses.

Characteristics of studies comparing MISC and OSC

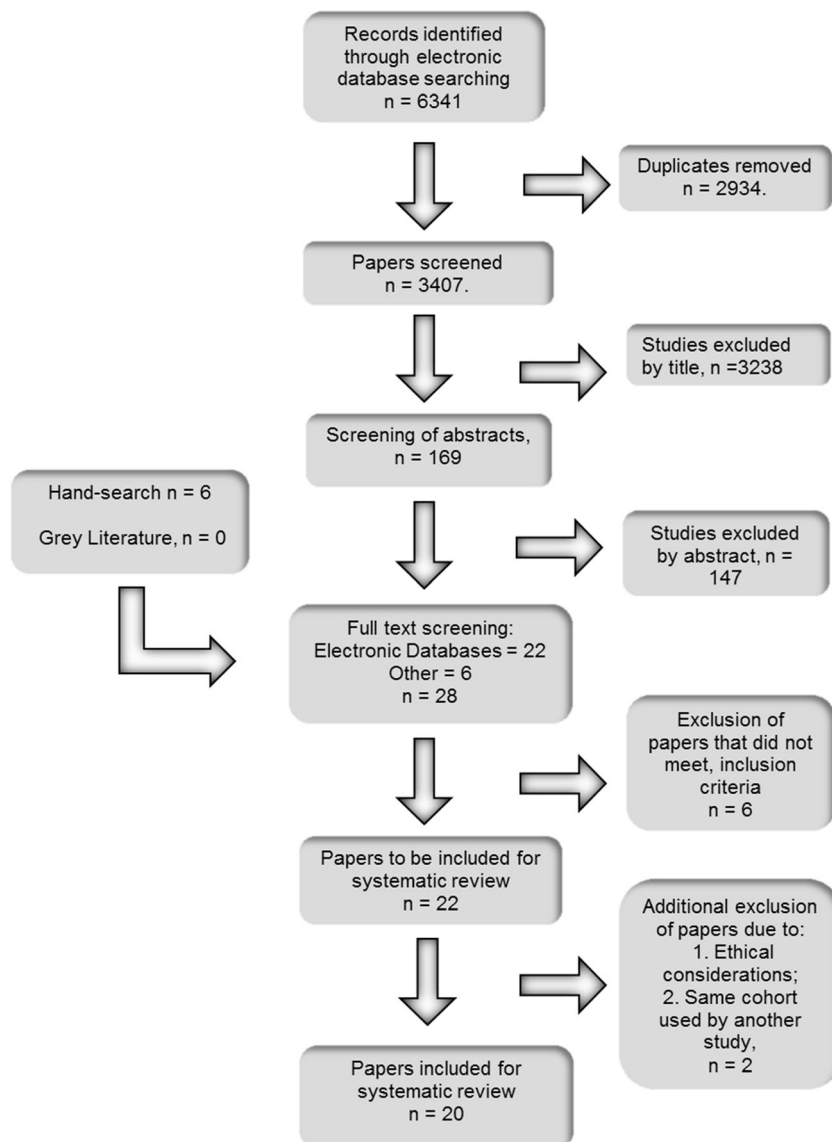
The characteristics of the studies comparing MISC and OSC are summarized in Table 3.

Study design

Of 12 studies, one was a RCT, two were prospective cohort studies and nine were retrospective cohort studies with a total of 4,757 subjects (Table 3). Furthermore, five studies compared RSC and OSC, four compared LSC and OSC and three compared MISC (RSC plus LSC) and OSC.

Technique used

Overall, five ports were used for RSC [12, 18, 19], whereas four ports were used for LSC [8, 20, 21]. In nine studies a concomitant hysterectomy was performed

Fig. 1 PRISMA flow chart**Table 2** Excluded studies

Study	Reason for exclusion
Hachem et al. 2013	Evaluated malignant and benign disease. No specific information regarding sacrocolpopexy was provided
Hsiao et al. 2007	Case series
Judd et al. 2010	Analysed costs in a hypothetical cohort. Costs was not a parameter to be evaluated in this systematic review
Marshall et al. 2010	Paper presented as an oral poster. No full article is available online. No email address is available to contact the authors
Unger et al. 2014	Paper presented as an oral poster. No full article is available online. The first author was contacted via email to obtain full article; no response was obtained
White et al. 2009	Compared a different approach (single port) which was not the objective of this systematic review
Siddiqui, Geller and Visco, 2012	Robotic cohort used seemed to be the same as that used by Geller et al. [19] (the time-frame was similar and both studies were conducted in the same institutions)
[29]	Ethical concerns. Ethical approval not mentioned. The journal's instructions for authors does not require a statement on ethical approval. The author was contacted but no response was obtained

Table 3 Included studies comparing MISC and OSC

Reference	Country	Comparison	Study design	Setting	Population (n)
[18]	USA	RSC vs. OSC	PC	Single centre	52 (RSC 30, OSC 22)
[20]	TheNetherlands	LSC vs. OSC	PC	Multicentre (four institutions)	85 (LSC 43, OSC 42)
[22]	USA	RSC vs. OSC	RC	Single centre	59 (RSC 40, OSC 19)
[27]	UK	LSC vs. OSC	RCT	Multicentre (not specified)	53 (LSC 26, OSC 27)
[19]	USA	RSC vs. OSC	RC	Multicentre (two institutions)	178 (RSC 73, OSC 105)
[23]	USA	RSC vs. OSC	RC	Single centre	84 (RSC 31, OSC 53)
[24]	USA	RSC vs. OSC	RC	Single centre	164 (RSC 73, OSC 91)
[25]	USA	MISC vs. OSC	RC	Multicentre (not specified)	970 (MISC 176, OSC 794)
[21]	USA	LSC vs. OSC	RC	Single centre	85 (LSC 44, RSC 41)
[12]	USA	MISC vs. OSC	RC	Multicentre (four institutions)	1,124 (MISC 535, OSC 589)
[28]	USA	LSC vs. OSC	RC	Single centre	117 (LSC 56, OSC 61)
[26]	USA	MISC vs. OSC	RC	Multicentre (about 200 hospitals)	1,786 (MISC 1,127, OSC 659)

RCT randomized controlled trial, RC retrospective cohort study, PC prospective cohort study

[12, 18, 19, 21–26]. Seven studies [12, 20, 21, 23, 24, 26, 27] reported the involvement of experienced surgeons, but in three studies [19, 22, 28] some surgeons were in their learning curve.

Quality appraisal

Figure 2 summarizes the risk of bias. Cohort studies were further appraised using the NOS (Table 4). Three studies [20, 24, 28] scored five stars, seven [12, 19, 21–23, 25, 26] scored six stars and one [18] scored seven stars.

Selection bias

One study [27] randomized subjects appropriately but failed to use adequate methods of allocation concealment. Inclusion of nonrandomized studies resulted in a high risk of selection bias (Fig. 3). Nevertheless, almost all cohort studies had a NOS score of three stars in the category “selection” (Table 4).

Performance bias

Only in one study [27] was the ward staff who supervised analgesic requirements blinded. It was not possible to blind subjects to the procedure because of the abdominal incision; however, this is unlikely to have introduced bias (Fig. 3).

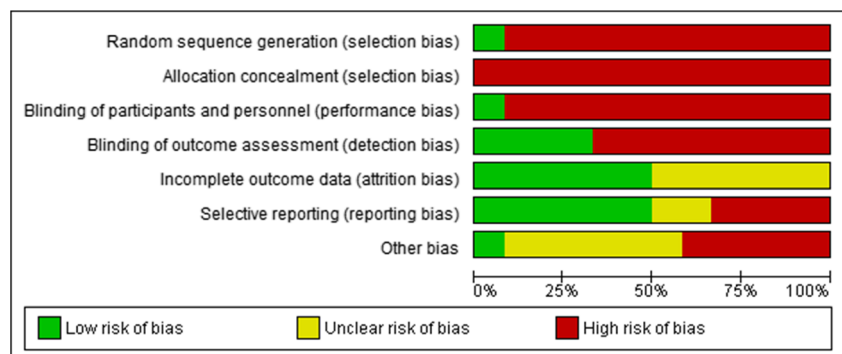
Detection bias

Blinding of outcome appraisal was performed in four studies [18, 23, 24, 27] (Fig. 3). Nonetheless, the NOS scores (Table 4) showed that almost all included cohort studies depended on patient records for assessment of outcomes.

Attrition bias

In six studies [18, 19, 22–24, 26], the follow-up rates were 80 % or more (Table 4, follow-up adequacy). Also, these studies had complete data, missing data were balanced across groups or were imputed using appropriate methods.

Fig. 2 Risk of bias: judgements of the review author about each risk of bias component displayed as percentages across studies comparing MISC and OSC



	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of participants and personnel (performance bias)	Blinding of outcome assessment (detection bias)	Incomplete outcome data (attrition bias)	Selective reporting (reporting bias)	Other bias
Collins et al., 2012	-	-	-	+	+	+	?
Coolen et al., 2013	-	-	-	-	?	-	?
Elliot et al., 2012	-	-	-	-	+	+	?
Freeman et al., 2013	+	-	+	+	?	+	-
Geller et al., 2008	-	-	-	-	+	+	-
Geller et al., 2012	-	-	-	+	+	-	-
Hoyte et al., 2012	-	-	-	+	+	+	-
Khan et al., 2013	-	-	-	-	?	?	-
Klauschie et al., 2009	-	-	-	-	?	+	?
Nosti et al., 2014	-	-	-	-	?	-	+
Paraiso, et al. 2005	-	-	-	-	?	-	?
Tyson and Wolter, 2013	-	-	-	-	+	?	?

Fig. 3 Risk of bias summary: judgement of review authors about each risk of bias component for each included paper comparing MISC and OSC

Reporting bias

Primary and secondary prespecified outcomes were reported in six papers [18, 19, 21, 22, 24, 27] (Fig. 3).

Other bias

One study [12] showed a low risk of bias due to funding and a conflict of interest, whereas five studies [19, 23–25, 27] showed a high risk of bias because of the source of funding or the presence of declared conflicts of interest (Fig. 3).

Confounders

Baseline characteristics able to act as confounders (age, body mass index, concurrent hysterectomy, other concurrent procedure or previous abdominal surgery) did not differ between groups in four studies [18, 20–22], and four studies [12, 23, 25, 26] used regression models to control for confounders.

Follow-up time

In six studies [10, 21, 23, 25, 27, 28], subjects were followed up for 12 months or more (Table 4, follow-up length).

Effectiveness of sacropexy for apical prolapse

Only four studies [19, 21, 23, 27] compared outcomes related to postoperative POP-Q assessments and cure rate (Table 5). Three studies [21, 23, 27] showed no significant differences in point-C POP-Q measurements 1 year after MISC or OSC. Freeman et al. [27] also found no significant difference in the satisfaction rate between MISC and OSC. Nosti et al. [12] found no significant difference in the apical prolapse recurrence rate at 1 year after controlling for confounding factors. These results are in accordance with those of other studies [21, 23, 25, 27, 28] (Table 5).

Complications of sacropexy

Nine studies compared complication rates between interventions [12, 19–21, 23, 25–28]. Complications were divided into intraoperative and postoperative. Events were extracted and included in the meta-analysis. Conversion to the open approach was not considered a complication. Two studies [12, 25] were not included in the meta-analysis because adverse events were reported as the sum of all complications. Overall complications (i.e. intraoperative plus postoperative) were similar between groups when combined in the meta-analysis (OR 0.91, 95 % CI 0.51 – 1.62, $p=0.74$; Fig. 4). However, Nosti et al. [12] found a higher rate of overall complications with OSC ($p<0.01$).

Intraoperative complications

Seven studies compared intraoperative complication rates between MISC and OSC (Fig. 5). The rate was 1 % for MISC and 2.4 % for OSC (Table 6). Although this favoured MISC, the difference was not statistically significant (OR 0.83, 95 % CI 0.51 – 1.34, $p=0.44$). This is comparable to previously reported results [25] showing no significant differences between groups ($p=0.1014$). Similarly, no significant differences were found when reported intraoperative complications were separately studied, including haemorrhage, bladder injury, intestinal injury

Table 4 Newcastle – Ottawa Quality Assessment Scale for each cohort study comparing MISC versus OSC

Newcastle – Ottawa Quality Assessment Scale. Cohort studies									
Study	Selection				Comparability	Outcome			
	Representativeness of the studied cohort	Selection of control cohort	Ascertainment of Exposure	Outcome at start of study	Comparability	Assessment	Follow-up length	Follow-up Adequacy	Total stars (out of 9)
Collins et al., 2012	b = ★	a = ★	b = ★	a = ★	a = ★	b = ★	b = 0	b = ★	7
Coolen et al., 2013	b = ★	b = 0	b = ★	a = ★	a = ★	c = 0	a = ★	d = 0	5
Elliot et al., 2012	b = ★	a = ★	a = ★	b = 0	a = ★	b = ★	b = 0	a = ★	6
Geller et al., 2008	a = ★	a = ★	a = ★	b = 0	b = ★	b = ★	b = 0	a = ★	6
Geller et al., 2012	b = ★	a = ★	a + b = ★	b = 0	none = 0	a + b = ★	a = ★	b = ★	6
Hoyte et al., 2012	b = ★	a = ★	a = ★	b = 0	none = 0	b = ★	b = 0	b = ★	5
Khan et al., 2013	b = ★	a = ★	a = ★	b = 0	b = ★	b = ★	a = ★	d = 0	6
Klauschie et al., 2009	b = ★	a = ★	a = ★	b = 0	a = ★	b = ★	a = ★	d = 0	6
Nosti et al., 2014	a = ★	a = ★	a = ★	b = 0	a + b = ★ ★	b = ★	b = 0	c = 0	6
Paraiso et al., 2005	b = ★	a = ★	a = ★	b = 0	none = 0	b = ★	a = ★	c = 0	5
Tyson and Wolter, 2013	a = ★	a = ★	a = ★	b = 0	b = ★	b = ★	b = 0	a = ★	6

and colpotomy (Appendices S1–S5). However, the transfusion rate was significantly lower with MISC (OR 0.41, 95 % CI 0.20 – 0.83, $p=0.01$; Fig. 6).

Postoperative complications

Of nine studies for which postoperative complications were reported, seven were pooled in the meta-analysis (Fig. 7). The rate was 5.41 % for MISC and 10.02 % for OSC (Table 6). However, no significant differences were identified (OR 0.82, 95 % CI 0.48 – 1.42, $p=0.64$).

No significant differences were found when postoperative complications were studied separately, including wound complications (disruption, infection, haematoma and hernia), fever, sepsis, thromboembolic events (deep venous thrombosis and pulmonary embolus), ileus/small-bowel obstruction/constipation, urinary retention, urinary

infection, urinary incontinence, abdominal pain requiring analgesics, pulmonary complications (pneumonia, unplanned reintubation and failure to wean ventilation within 48 h), neurological complications (cerebrovascular accident, peripheral nerve injury and delirium) (Appendices S6–S16), and mesh erosion (Fig. 8). Similarly, Nosti et al. [12] found no significant differences in thromboembolic events, wound complications and mesh erosion.

Length of hospital stay

Of nine studies [12, 19–22, 24, 26–28] for which LOS was reported, only four could be combined in the meta-analysis (Fig. 9). LOS (expressed in days) was significantly shorter following MISC (MD -1.57 days, 95 % CI -1.91 – -1.23 days, $p<0.00001$; Fig. 9). These results are in

Table 5 Characteristics of studies comparing MISC with OSC

Reference	Study design	Groups compared	Population (n)	Procedure	Follow-up time	Effectiveness ^a				Apical prolapse recurrence	Surgeon's experience	
						1 month	3 months	6 months	1 year			2 years
[19]	RC	RSC vs. OSC	178 (RSC 73, OSC 105)	Sacrocolpopexy, sacrocervicopexy	6 weeks	SD (<i>p</i> =0.008)	NR	NR	NR	NR	NR	Some in learning curve
[23]	RC	RSC vs. OSC	84 (RSC 31, OSC 53)	Sacrocolpopexy, sacrocervicopexy	44 months	NR	NR	NR	NR	NR	NR	Experienced
[22]	RC	RSC vs. OSC	59 (RSC 40, OSC 19)	Sacrocolpopexy, sacrocervicopexy	30 days	NR	NR	NR	NR	NR	NR	Some in learning curve
[24]	RC	RSC vs. OSC	164 (RSC 73, OSC 91)	Sacrocolpopexy, sacrocervicopexy	3 and 6 months	NR	NR	NR	NR	NR	NR	Experienced
[18]	PC	RSC vs. OSC	52 (RSC 30, OSC 18)	Sacrocolpopexy, sacrocervicopexy	10 days	NR	NR	NR	NR	NR	NR	NM
[28]	RC	LSC vs. OSC	117 (LSC 56, OSC 61)	Sacrocolpopexy	OSC 15.7±18.1 months, RSC 13.5±12.1 months	NR	NR	NR	NR	NR	NR	1 case in LSC group (<i>p</i> =NR)
[21]	RC	LSC vs. OSC	85 (LSC 44, OSC 41)	Sacrocolpopexy, sacrocervicopexy	OSC 7.4 months, LSC 10.6 months	SD (<i>p</i> =0.002)	NR	NR	SD (<i>p</i> =0.009)	NR	NR	No recurrence (<i>p</i> =NR)
[20]	PC	LSC vs. OSC	85 (OSC 42, LSC 43)	Sacrocolpopexy	1 year	NR	NR	NR	NR	NR	NR	Experienced
[27]	RCT	LSC vs. OSC	53 (OSC 27, LSC 26)	Sacrocolpopexy	3 months and 1 year	NR	NSD (<i>p</i> =NR)	NR	NSD (<i>p</i> =NR)	NR	NR	Experienced
[12]	RC	MISC vs. OSC	1,124 (OSC 589, MISC 535=LSC 273, RSC 262)	Sacrocolpopexy, sacrocervicopexy	LSC 8 months, OSC 8 months, RSC 14 months	NR	NR	NR	NR	NR	NR	Experienced
[25]	RC	MISC vs. OSC	970 (OSC 794, MISC 176)	Sacrocolpopexy, sacrocervicopexy	1 year	NR	NR	NR	NR	NR	NSD (<i>p</i> =NR)	NM
[26]	RC	MISC vs. OSC	1,786 (OSC 659, MISC 1,127)	Sacrocolpopexy, sacrocervicopexy	30 days	NR	NR	NR	NR	NR	NR	NM

NM not mentioned, NR not reported, SD significant difference, NSD not significant difference, RC retrospective cohort study, PC prospective cohort study, RCT randomized controlled trial

^a Objective apical cure rate (point C POP-Q, stage ≤1). *p* values indicate the significance of differences between groups

Fig. 4 Comparison of overall complications between MISC and OSC

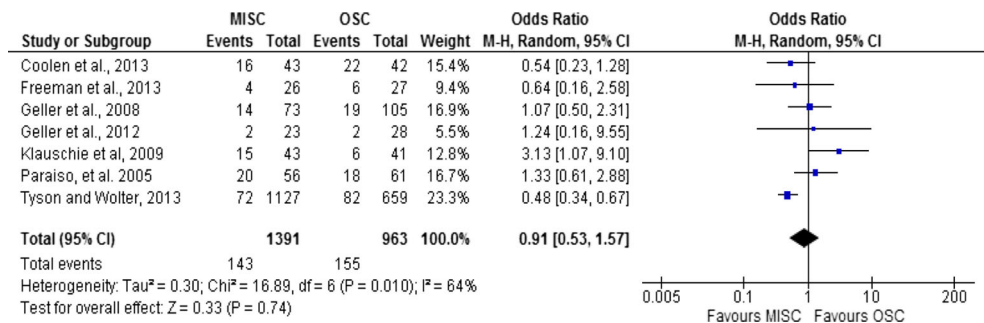


Fig. 5 Comparison of overall intraoperative complications between MISC and OSC

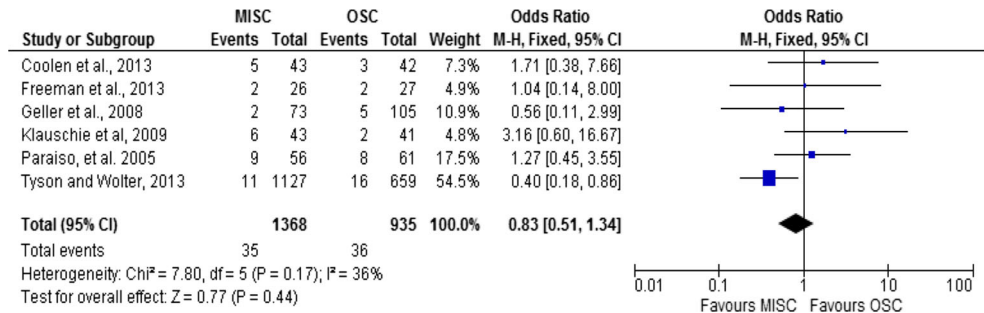


Table 6 Complication rates of MISC and OSC

Complication	Complication rate (%)	
	MISC	OSC
Intraoperative	2.55	3.85
Haemorrhage	2.36	1.75
Bladder injury	0.87	0.64
Bowel injury	0.07	0.42
Postoperative	7.76	12.04
Overall	10.31	15.89

accordance with those of other studies [20–22, 24, 26] (all $p < 0.001$).

Estimated blood loss

Of nine studies [12, 19–22, 24, 27, 28] for which EBL was reported, only five were combined in the meta-analysis

(Fig. 10). EBL was significantly greater with OSC than with MISC (MD -113.27 mL, 95 % CI -163.67 – -62.87 mL, $p < 0.0001$; Fig. 10). These results are in accordance with those of other studies [20–22, 24].

Operating time

OT was evaluated in ten studies [12, 18–22, 24, 26–28]. OT was defined differently among the studies so inclusion in the meta-analysis resulted in substantial heterogeneity. Consequently, a subgroup analysis was performed. “Room operating time” (i.e. total time in the operating theatre) was shorter with OSC [24, 28]. Similarly, “from incision to closure” OT was shorter with OSC in five studies [12, 19, 21, 24, 26]. Meta-analysis of two studies [12, 19] showed a significantly shorter OT with OSC (MD 87.47 min, 95 % CI 58.60 – 116.34 min, $p < 0.0001$; Fig. 11). In four studies [18, 20, 22, 27], OT was not specifically defined.

Fig. 6 Comparison of transfusion rates between MISC and OSC

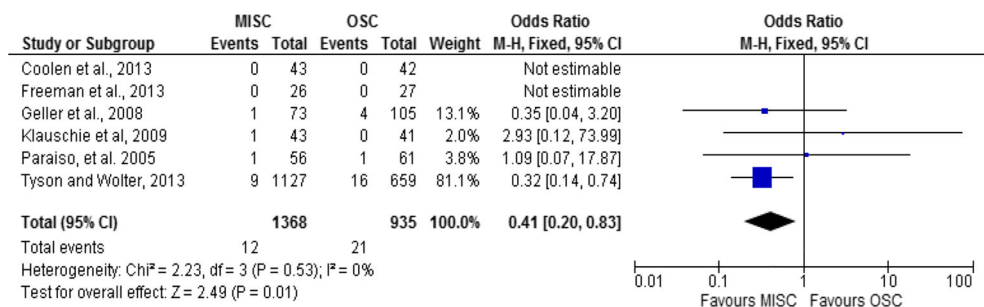


Fig. 7 Comparison of overall postoperative complications between MISC and OSC

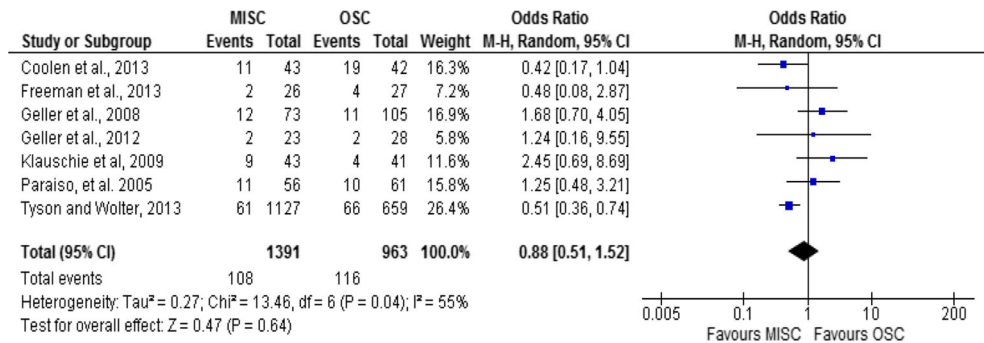


Fig. 8 Comparison of mesh erosion rates between MISC and OSC

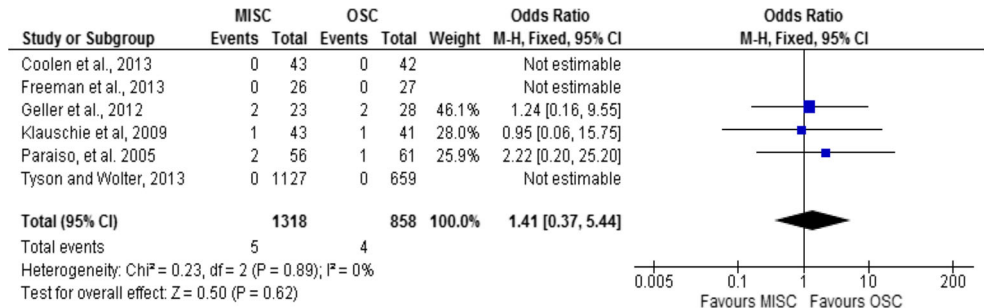
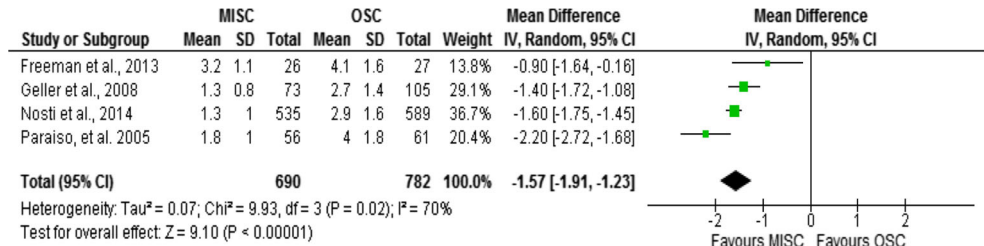


Fig. 9 Comparison of length of stay (days) between MISC and OSC



Interestingly, none of them showed significant differences between procedures. However, in one study [22] this was attributed to the fact that OSC was probably used to treat more complex cases. Hoyte et al. [24] also evaluated the OT required only for sacropexy and found shorter OT with OSC ($p < 0.001$).

Postoperative pain

Postoperative pain was evaluated in three studies. Freeman et al. [27] evaluated morphine use over 3 days, whereas Collins et al. [18] measured the number of postoperative oral narcotics required. Although usage of postoperative

Fig. 10 Comparison of estimated blood loss (millilitres) between MISC and OSC

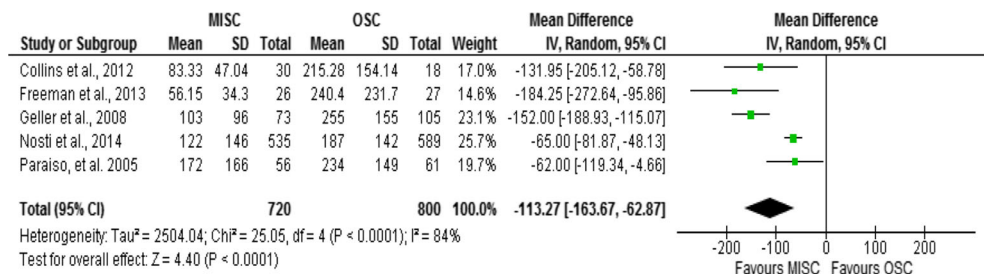
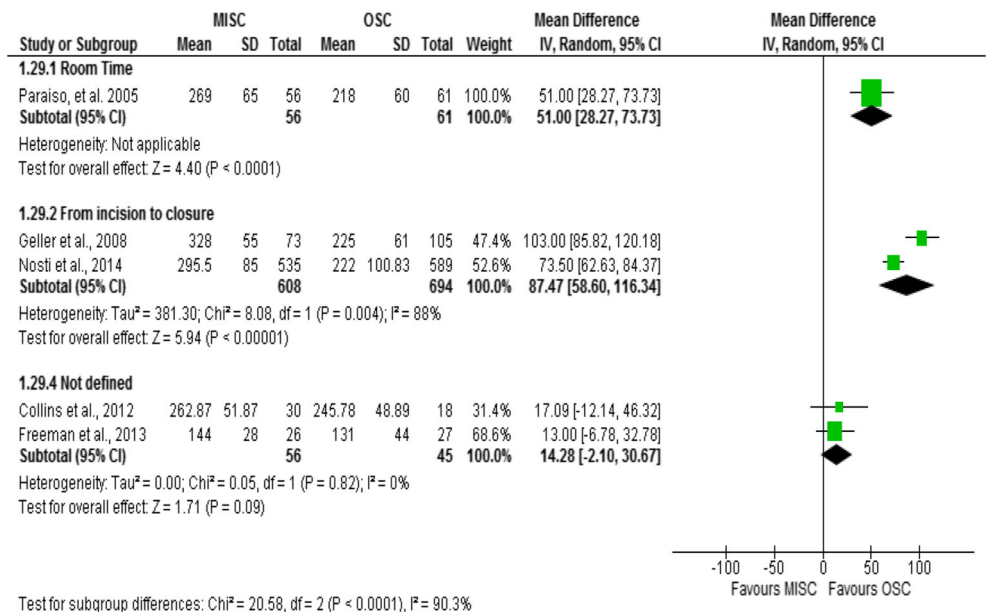


Fig. 11 Comparison of operating times (minutes) between MISC and OSC



analgesics appeared to be less with MISC, the difference did not reach statistical significance (SMD -0.41, 95 % CI -0.83 – 0.01, *p*=0.05; Fig. 12). Similarly, Khan et al. [25] found no significant differences in perioperative pain between groups (*p*=0.14).

Postoperative quality of life

Postoperative QoL was evaluated in three studies [18, 23, 27]. Two studies [18, 27] showed no significant differences between groups applying the Short Form-36 Health Survey. Similarly, applying the Prolapse QoL Questionnaire, one study [27] showed no significant differences between the approaches (*p*=0.95). Additionally, no significant differences were found for the subscales of the PFDI-20 and PFIQ-7 questionnaires (*p*=0.68) or the PISQ-12 questionnaire (*p*=0.32) [23].

Mortality

In two studies [20, 26] fatalities occurred after MISC and OSC. In one study [20] one patient died in the open arm due to sepsis and subsequent multiorgan failure after a bowel

perforation, and in the other [26] one patient died in each group, but the causes of death were not specified. When combined in the meta-analysis no significant differences were found (OR 0.44, 95 % CI 0.06 – 3.48, *p*=0.44; Fig. 13).

Secondary analyses

Publication bias

Funnel plots were used to find the risk of publication bias (Appendices S17–S23). However, in cases of funnel plot asymmetry, further tests could not be done because fewer than ten studies were included in the meta-analysis for each outcome [9].

Sensitivity analyses

After removing observational studies with a low NOS score, the pooled estimate of effect remained similar between groups for all outcomes studied. Only the results related to “complications of sacropexy” would have changed if fixed-effects analysis had been performed instead. However, this would have placed excessive weight on one observational study [26] reducing the weight of the more methodologically

Fig. 12 Comparison of postoperative pain between MISC and OSC

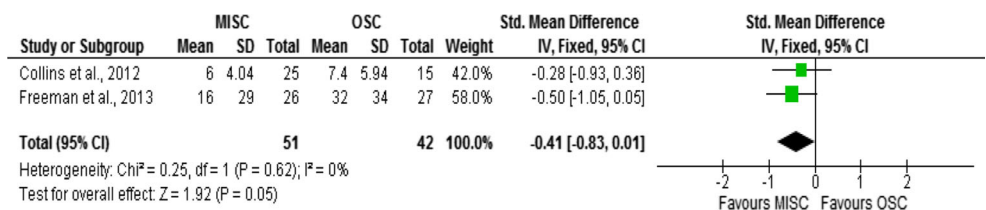
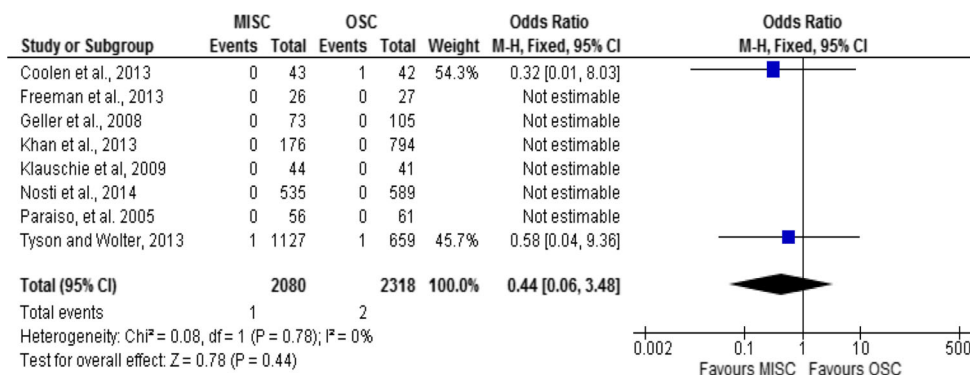


Fig. 13 Comparison of mortality between MISC and OSC

rigorous study [27]. Studies with participation of experienced surgeons and surgeons in their learning curve were also compared. Studies in which concomitant hysterectomy was performed were also compared with those in which hysterectomy was not done. These comparisons revealed no differences between the surgical approaches, indicating that the results obtained are highly robust.

Discussion

Of the previous systematic reviews on this topic, the meta-analysis by Reza et al. [7] and the Cochrane review [8] only included one study each on sacrocolpopexy [19, 28], which were included in the present systematic review. Consequently, they are not discussed further. Additional consideration was given to the systematic review by Serati et al. [10]

Main findings

Twelve studies involving 4,757 participants compared MISC and OSC. The quality of the included studies was variable. Eleven studies were nonrandomized, and confounders were present in the majority. However, only three nonrandomized studies [20, 24, 28] had a low NOS score (five stars). It is clear though that poor methodological rigour in the included studies may have reduced the strength of the results, and cautious interpretation is advised. At 1 year postoperatively, MISC and OSC were equally effective in terms of point-C POP-Q measurements and recurrence rate. While mortality rate and the overall, intraoperative and postoperative complication rate favoured MISC, these differences did not reach statistical significance. Specifically, no significant differences were found in the mesh erosion rate. QoL appeared to be similar between MISC and OSC in the different questionnaires evaluated. MISC was also associated with less postoperative pain in terms of the quantity of postoperative analgesics. However, the difference was not statistically significant.

Strengths and limitations

Some weaknesses of this review must be addressed. Selection bias was inevitable due to the inclusion of nonrandomized studies. However, the scarcity of RCTs comparing MISC and OSC necessitated the inclusion of nonrandomized studies. Additionally, although articles in English, Portuguese and Spanish were sought, language bias is also present. This may have reduced the precision of the summary effect in the data analysis.

Many reasons for heterogeneity were detected: i.e. experience of surgeons, inclusion of concomitant hysterectomy and the differences in the techniques used among the studies. Specifically, regarding the outcome complication rate, performing hysterectomy at the time of promontofixation and the experience of surgeons were found to have an important influence on the results. Consequently, it could be argued that studies in which hysterectomy was concomitantly performed should have been excluded from this review. Nonetheless, it is believed that the trend seen among the studies is common in daily practice and therefore this review reflects real-world results.

Random-effects meta-analyses were performed when heterogeneity was high. Arguably, using analyses can exacerbate the risk of bias of smaller studies [9]. However, studies with larger populations were also less methodologically rigorous. Using random-effects analyses allowed the average of the effects to be obtained balancing the results more appropriately and giving more weight to studies with less risk of bias. Finally, it would have been valuable to explore results regarding the anterior and posterior compartments, postoperative stress urinary incontinence, costs and conversion rate between approaches, but this would have made the review unmanageable.

Despite these limitations, we believe that our results are valid. The comprehensive search used, the inclusion of trial registries in the search strategy and the attempts

made to include unpublished studies would have reduced the risk of publication bias. Additionally, sensitivity analyses showed consistent results in the majority of outcomes. Furthermore, each step of the review was undertaken following a systematic approach based on the recommendations of The Cochrane Handbook [9] using a predefined protocol with strict inclusion and exclusion criteria as well as a thorough critical appraisal using appropriate assessment tools.

Interpretation

Regarding the effectiveness of MISC versus OSC, our results are in accordance with those found in a literature review [30] comparing LSC and OSC. A previous systematic review comparing RSC and OSC did not provide figures regarding anatomic outcomes but concluded that the two procedures were comparable [10]. The inclusion of a RCT in our review reinforces the existing assumptions showing clinical equivalence between MISC and OSC.

There was high heterogeneity in the overall complication rate. The causes were thought to be the varied experience of surgeons and the inclusion of concomitant hysterectomy, but sensitivity analyses do not support this. Nonetheless, one outlier was detected. The study by Klauschie et al. [21] showed a higher rate of complications with MISC. If this study was excluded from the group of experienced surgeons, MISC showed a significantly lower complication rate than OSC ($p < 0.00001$). This may suggest that the complication rate could be lower with MISC if experienced surgeons perform the procedure. A meta-analysis [31] comparing laparoscopy with laparotomy in benign gynaecological pathology showed a significantly lower overall complication rate with laparoscopy; however, the learning curve was not considered in that analysis. It is not clear why the complication rate reported by Klauschie et al. [21] differed from that in other similar studies. Claerhout et al. [32] found that satisfactory learning in LSC is achieved after 60 cases. None of the papers except two [20, 27] reported the number of procedures performed by surgeons. Future clarification in this regard may help to explore the influence of the learning curve on this and other outcomes.

Similar to the findings of Barber and Maher [30], this systematic review found that MISC was superior to OSC in terms of transfusion rate, LOS and EBL. Again, high heterogeneity was found in LOS and EBL. Reasons that might have influenced these outcomes are the number of other additional procedures performed concomitantly at the time of sacropexy, differences in population characteristics, differences among institutions in their policy on postoperative stay, preoperative stage of prolapse, and complexity of cases. Shorter LOS with laparoscopy was also found in a Cochrane review [33] of benign ovarian tumour and

the review by Serati et al. [10] who also found shorter LOS and less EBL with RSC than with the open approach.

Similar to the findings of Barber and Maher [30], conflicting results were obtained regarding OT. Thorough analysis was not possible because clear definitions of OT were not provided for four studies. Additionally, the type of procedure performed, the preoperative stage of prolapse, and the varied findings during procedures possibly contributed to the heterogeneity observed. However, baseline characteristics in three studies [12, 24, 26] showed significantly more prior abdominal surgery in the open arm. This would have increased OT (due to potentially more adhesions). Nevertheless, OT in such studies were still shorter with OSC. Consequently, shorter OT was associated with OSC, as also found by Serati et al. [10]. Conversely, MISC showed less postoperative pain in terms of the quantity of postoperative analgesics. Although this difference did not reach statistical significance, only two studies were available. More research regarding postoperative pain may confirm a statistically significant difference between surgical techniques for sacropexy similar to the findings in other conditions such as benign ovarian tumours [33] and endometrial cancer [34] in which postoperative pain is less with the laparoscopic approach.

Conclusions

Practical recommendations

Current evidence regarding sacropexy for the treatment of prolapse of the apical segment of the vagina indicates comparable anatomic outcomes between MISC and OSC at 1 year postoperatively. MISC was associated with a lower transfusion rate, shorter LOS and less EBL but also with longer OT. The rate of other complications was comparable between the approaches. MISC shows benefits over OSC and can be considered in clinical settings where technology and surgical proficiency are accessible. Cautious interpretation of the results is advised due to the high risk of bias.

Research recommendations

It is paramount to emphasize the importance of further high-quality research (RCTs) in order to minimize bias and reduce the effects of confounders. Similarly, precise definitions are required in future trials. The lack of clear concepts of OT limited the depth of the analysis. Furthermore, “surgeon’s expertise” may be defined to explore more accurately the influence of the learning curve on different outcomes. Alternatively, future studies may provide the approximate number of procedures that have been performed per approach by surgeons. Moreover, a supplementary

systematic review evaluating results regarding the anterior and posterior compartments, postoperative stress urinary incontinence, costs, conversion rate and ergonomics will support the decision as to the best approach to sacropexy.

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