Is Single Incision Laparoscopic Cholecystectomy Really Less Invasive than Traditional Laparoscopic Cholecystectomy

Vivek Srivastava*, Narendra Nath Das**, Anand Kumar***, Mumtaz Ahmad Ansari***

*Assistant Professor, **Junir Resident, ***Professor, Department of Surgery, Institute of Medical Sciences, Banaras Hindu University, Varanasi, Uttar Pradesh, India.

Abstract

Objective: Our objective was to assess the clinical outcomes (benefits and drawbacks) and change in level of inflammatory parameter IL-6 in patients undergoing single incision laparoscopic cholecystectomy and comparison with classical four port laparoscopic cholecystectomy. Methods: Between September 2013 to July 2015 a prospective randomised study was conducted. Sixty patients were included in the study and they underwent elective gall bladder removal by applying the laparoscopic technique. All the patients were divided into two groups. Single incision laparoscopic cholecystectomy (group I) and four port laparoscopic cholecystectomy (group II). Outcome Measures included operative time, pain intensity post operatively and consumption of pain killers, hospital stay, need for conversion, complications, cosmetic effects and change in the level of serum Interleukin-6 post operatively as an inflammatory marker. Results: Mean operating time in group I was 71 min and group II 39 min. Intensity of pain evaluated by using the VAS at 8 hours after surgery in group I was 6.5 and in group II 6.5, whereas after 7 days in group I it was 2.7 and in group II 3.6. The pain killer requirement in group I was smaller than group II. Mean hospital stay after the operation in group I was 2.2 days and in group II 2.0 days. There were 2 conversions in group I and 1 in group II. Cosmesis evaluated by a 0 to 10 scoring system which showed better cosmesis in group I patients. Change in the serum level of IL-6 post operatively was more in case of multiport laparoscopic choecystectomy than single incision laparoscopic cholecystectomy. Single incision Conclusion: laparoscopic

E-mail: mumtazbhu@gmail.com

cholecystectomy is a safe and feasible procedure which has a better cosmesis and faster recovery.

Keywords: Laparoscopic Cholecystectomy (LC); Symptomatic Gallstone Disease; Single Incision Laparoscopic Cholecystectomy (SILC).

Introduction

Laparoscopic cholecystectomy (LC) is the gold standard treatment for benign and symptomatic gallstone disease [1,2]. Its main advantages over open cholecystectomy are the reduced early post-operative pain, shorter hospital stay, rapid return to the normal activity and better cosmesis. The continuous endeavour to reduce the invasiveness and thus the reduction of wound related complications and betterment of cosmesis following surgery has led the surgeons to further reduce the number and size of access ports during laparoscopic procedure. Various natural orifices including the trans-gastric, transrectal, and trans-vaginal route have been used as access although limited by lack of reproducibility, longer learning curve and ethical issues [3-6]. To reduce the invasiveness of standard four port cholecystectomy, single incision laparoscopic cholecystectomy (SILC) has also become an attractive option of the performance of laparoscopic cholecystectomy [7-10]. Navarra et al first reported trans-umbilical single incision laparoscopic cholecystectomy in 1997 and proposed that SILC might be associated with less pain and reduced hospitalization [11]. However, there was not enough data to support SILC as the standard of care as compared to multiport laparoscopic cholecystectomy as it was associated with longer operating time and required additional instruments more frequently. Inteleukin-6 (IL-6) is an inflammatory marker assessed post operatively which can be used as a surrogate marker of inflammation and can predict

Corresponding Author: Mumtaz Ahmad Ansari, Professor, Department of Surgery, Institute of Medical Sciences, Banaras Hindu University, Varanasi, Uttar Pradesh 221005, India.

the overall surgical stress, an important factor for recovery. There are many studies comparing laparoscopic cholecystectomy and open cholecystectomy reported significantly lower level of IL-6 post-operatively in the laparoscopic groups, which suggests that the minimally invasive approach is less stressful. There are very few studies comparing the rise of IL-6 post operatively between SILC with conventional laparoscopic cholecystectomy and the results are inconclusive also.

Hence, we planned a study to compare the various outcome parameters and level of rise of IL-6 between SILC and traditional four port LC.

Material and Methods

The study was conducted at a University hospital between September 2013 to July 2015 among consecutive patients undergoing elective LC for ultrasonographically diagnosed symptomatic gallstone disease aged between 18-70 years with ASA I/II score. The exclusion criteria were suspected Mirrizi syndrome, common duct stones or malignancy, deranged coagulation profile and with acute cholecystitis or choledocholithiasis proven on ultrasound. After enrollment in the study the patients were randomly allocated to Single incision Laparoscopic Cholecystectomy (SILC) group, Three port LC (3LC) group or Conventional four port LC (4LC) group by use of computer generated random number table.

Anaesthesia Technique

The anaesthesia techniques, anaesthetic drugs and surgical techniques were standardized. Anesthesia was induced with propofol 2–2.5 mg/kg IV, glycopyrrolate 0.2 mg IV, and fentanyl 2 ig/kg IV. Endotracheal intubation was facilitated with vecuronium 0.15 mg/kg IV. Anesthesia was maintained with 1.0%–2.5% (inspired concentration) isoflurane in oxygen. Ventilation was controlled mechanically and adjusted to keep an end-tidal CO2 partial pressure of 30–40 mm Hg. Neuromuscular block was maintained with vecuronium IV. After tracheal intubation, a nasogastric tube was placed to promote baseline emptying of the stomach of air and gastric contents.

Surgical Technique for Single Incision Laparoscopic Cholecystectomy (SILC)

Patient was placed in reverse trendelenburg leg

apart position with the table tilted downward to the patient's left (15 degree). After infiltration of 5 ml bupivacaine solution around the umbilicus, transumbilical 2.5-3.5 cm incision was given and SILC port was placed in the umbilicus through the same skin incision. Pneumoperitoneum was created with insufflation of the abdomen with C02 using SILS port inlet at 15 mm Hg. The two 5mm trocars were used for introducing the convention LC hand instruments. The central 5 mm port was used to introduce 30 degree long telescope (50 cm). An initial gross examination of the entire abdomen cavity was made primarily to exclude injury/bleeding during the creation of the pneumoperitoneum. The anatomy was visualized. A Maryland dissector and a grasper were introduced through other 5mm trocars introduced in SILS port. The fundus of the gallbladder was grasped initially and progressing gradually to Hartman's pouch doing necessary adhesiolysis and flipped upwards over the superior edge of the right lobe by a curved grasper through the SILC port. The sufficient length of cystic duct and cystic artery on gallbladder side were skeletonized, clipped with liga-clips and divided making sure to visualize the gallbladder cystic duct junction and common bile duct cystic duct junction wherever possible. The dissection of the gallbladder from the liver bed was done by laparoscopic hook with monopolar cautery. The gallbladder was then held with toothed grasper and brought out through the umbilical incision. Any bile spill was irrigated with normal saline and suctioned and any stone spill were retrieved. Rectus sheath at umbilicus was closed with vicryl no 1 and skin was approximated with stapler.

Surgical Technique for Standard 4 port Laparoscopic Cholecystectomy

All patients were placed in reverse Trendelenburg position with 15 degree left lateral tilt. Premptive analgesic with 5 ml bupivacaine solution was given at the site of incision and pneumoperitoneum was created via Veress needle with closed technique. A 10-mm supraumbilical port was placed for camera and 3 working ports were made- 10mm port in the mid-epigastrium just to the right of the falciform ligament, and two 5-mm ports in the right upper abdomen two finger breadths below the right margin in the mid-clavicular and the anterior-axillary line. A 10 mm 0° laparoscope was used. An initial gross examination of the entire abdomen cavity was made primarily to exclude injury/bleeding during the creation of the pneumoperitoneum and secondly to identify any gross macroscopic additional disease. The fundus of the gallbladder was grasped by the assistant and flipped upwards and over the superior edge of the right lobe. A Maryland dissector and a grasper was used to identify the structures in the Calot's triangle using monopolar cautery. The sufficient length of cystic duct and cystic artery on gallbladder side were skeletonized, clipped with 10 mm liga-clips and divided making sure to visualize the gallbladder cystic duct junction and common bile duct cystic duct junction wherever possible. The dissection of the gallbladder from the liver bed was done by laparoscopic hook with monopolar cautery. The gallbladder was then held with toothed grasper and brought out through the epigastric incision. Any bile spill was irrigated with normal saline and suctioned and any stone spill were retrieved.

Assessed Factors

In this study we assessed the conversion rate, duration of surgery, degree of postoperative pain, use of analgesics, hospital stay, complications, cosmetic satisfaction and change in serum level of interleukin 6 postoperatively.

Conversion was assessed by change from one surgical procedure to another for successful removal of gallbladder. The reasons for conversion were recorded. The port site wound infections were classified according to the CDC classification [12] for surgery site infections. Operative time was measured in minutes defined as time taken from, start of giving first incision to skin closure of the last incision. Severity of postoperative pain was recorded at 8hrs after operation and during follow up at 1 week, 3 months by using Visual Analogue Scale (VAS). Hospital stay was calculated as the number of days in the hospital after the surgery until the patient was deemed fit for the discharge by operating surgeon. Cosmetic satisfaction of surgical scar was rated on a scale [range, 0(worst) to 10(best)] and was evaluated at POD 7 or stitch removal which one is earlier and at the 3 month and 6 month follow up visit.

Pre operatively serum level of IL-6 was measured in all patients. Post operatively after 24 hr serum level of IL-6 was re-measured. IL-6 estimation was done by chemiluminesence immunoassay system kit. Then level of increase of this inflammatory mediator was compared between the groups.

Statistical Analysis

Statistical analysis was done using SPSS version 16.0. For continuous data ANOVA test was used to compare the significant difference in mean for more than two groups. For categorical variables Chi-square test and Fisher's – exact test were used. The P-value of < 0.05 considered as statistically significant.

Results

Of 46 patients enrolled in the study 40 patients completed the study. One patient had incidentally diagnosed carcinoma gallbladder on laparoscopy, two had uncontrolled comorbidity, one patient had Mirrizi syndrome and two patient lost to followup after discharge. A total of 40 patients were randomized into two groups. Group I underwent SILC and Group II 4 port LC. The patient characteristics between the two groups were comparable (Table 1a and 1b). There were 2 conversions in Group and 1 in Group II (pvalue-0.804) all because of non progression due to dense Calot's adhesions. There was no difference in the post operative analgesic intake, type of analgesic used and the rate of surgical site wound infection rate (Table 2). The postoperative outcomes are shown in Table 3. The SILC group significantly longer operating time when compared to conventional LC group (Mean time 71 versus 39.5 minutes) although the duration of hospital stay was similar. The VAS was similar on day 0 but was significantly less in SILC group on day 7 and at 3 months postoperatively. Similarly, the day of resuming work was significantly shorter in SILC group compared to conventional LC. The rise in IL-6 level was significantly higher in conventional group compared to SILC group (p-value-<0.001). The cosmesis as assessed by the patient was

Table 1a: Comparison of preoperative parameters between the groups

| Variables | | SILC | | 4 port | | |
|-----------------------|--------|------|-------|--------|-------|---------|
| (N=40) | | No. | %. | No. | %. | p-value |
| Sex | Male | 3 | 15.0 | 1 | 5.0 | 0.418 |
| | Female | 17 | 85.0 | 19 | 95.0 | |
| Socio-economic Status | High | 3 | 15.0 | 2 | 10.0 | 0.767 |
| | Middle | 17 | 85.0 | 17 | 85.0 | |
| | Low | 0 | 0.0 | 1 | 5.0 | |
| Dyspepsia | Yes | 20 | 100.0 | 20 | 100.0 | 0.001 |
| | No | 0 | 0.0 | 0 | 0.0 | |
| Pain abdomen | Yes | 20 | 100.0 | 20 | 100.0 | 0.362 |
| | No | 0 | 0.0 | 0 | 0.0 | |

New Indian Journal of Surgery / Volume 6 Number 4 / October - December 2015

Vivek Srivastava et. al. / Is Single Incision Laparoscopic Cholecystectomy Really Less Invasive than Traditional Laparoscopic Cholecystectomy

| Acid Peptic Disorder | Yes No | 20 0 | 100.0 0.0 | 18 2 | 90.0 10.0 | 0.006 |
|----------------------|-----------|---------|--------------|---------|--------------|-------|
| Previous operation | Yes No | 2 18 | 10.0 90.0 | 1 19 | 5.0 95.0 | 0.804 |
| Diabetes Mellitus | Yes | 0 20 | .0 100.0 | 2 18 | 10.0 90.0 | 0.349 |
| Tenderness | Yes No | 1 19 | 5.0 95.0 | 0 20 | 0.0 100.0 | 0.362 |

Table 10b: Comparison of pre-operative parameters between the study groups

| Variables (N=40) | SILC (Mean±SD) | 4 port (Mean±SD) | F-value | P-value |
|---------------------|------------------|------------------|---------|---------|
| Age | 36.60±9.087 | 41.15±13.299 | .775 | .465 |
| Hemoglobin | 12.1350±1.30436 | 12.4150±0.96860 | .830 | .441 |
| Total Count | 8266.50±1481.792 | 7650.00±1612.615 | .830 | .441 |
| Creatinine | 0.8050±0.19861 | 0.6550±0.25021 | 2.474 | .093 |
| Urea | 25.125±8.7778 | 31.600±8.3376 | 2.956 | .060 |
| SGPT | 33.760±8.8448 | 48.200±20.0935 | 3.495 | .037 |
| SGOT | 35.775±7.7366 | 43.100±10.0990 | 1.945 | .152 |
| Direct Bilirubin | 0.260±0.1429 | 0.340±0.1314 | 1.950 | .152 |
| Total Bilirubin | 0.7200±0.24192 | 0.8450±0.16694 | 4.048 | .023 |
| Alkaline | 103.305±26.8524 | 110.700±18.4622 | .538 | .587 |
| Phosphatase | | | | |
| Total Protein | 7.8050±0.57626 | 7.7650±0.69606 | 6.733 | .002 |
| Albumin | 4.3550±0.56240 | 4.0050±0.47404 | 3.181 | .049 |

SILC 4 port P-value % % No No Failure of the technique (n=60) Yes 2 10.0 5.0 1 0.804 18 90.0 19 95.0 No Nature analgesic agents (n=60) **NSAIDs** 15 75.0 14 70.0 0.377 NSAIDs and 5 25.0 6 30.0 opioid 2 Wound infection (n=60) Yes 10.0 1 5.0 0.765 No 18 90.0 19 95.0

 Table 3: Comparison of post-operative parameters

| Variables | Group 1 Mean±SD | Group 3 Mean±SD | F-value | P-value | |
|----------------------------------|--------------------|--------------------|---------|---------|--|
| Duration surgery | 71.00±9.403 | 39.50±9.162 | 92.209 | < 0.001 | |
| Duration hospital stay | 2.20±0.523 | 2.00±0.459 | 1.541 | 0.223 | |
| Pain day 0 (VAS score) | 6.50±0.889 | 6.50±1.100 | 2.367 | 0.103 | |
| Pain day 7 (VAS score) | 2.70±0.979 | 3.60±1.046 | 18.455 | <0.001 | |
| Pain 3 month (VAS score) | 1.60±0.821 | 1.90±0.788 | 6.077 | 0.004 | |
| Resuming daily work POD | 6.65±1.182 | 5.75±1.070 | 5.443 | 0.007 | |
| IL6 preop | 15.3185±7.13412 | 21.5100±11.30472 | 1.060 | 0.353 | |
| IL6 postop | 76.15 ? 19.83 | 155.23±80.50240 | 12.04 | <0.001 | |
| Cosmetic satisfaction Day 7 | 7.20±1.361 | 5.20±1.196 | 16.964 | <0.001 | |
| Cosmetic satisfaction 3months | 9.30±1.342 | 7.20±1.196 | 31.683 | <0.001 | |
| Cosmetic satisfaction 6months | 9.60±0.821 | 7.80±0.894 | 34.977 | <0.001 | |

significantly better in SILC group at day 7 and 3 and 6 months after the operation.

Discussion

134

disease. The technique of LC has been standardised and the outcome of the patients following LC is almost stable with a conversion rate of 0.2% [13], biliary complication rate of 0.26 to 0.6% [14,15] and bowel injury rate of 0.14 to 0.35 % [14,15]. Majority of the morbidity related to pain, wound complications and cosmetic outcomes are related to the access sites for

LC is the gold standard treatment for gall stone

LC. There has been a continuous endeavour to reduce the invasiveness and thus wound related complications of LC and also improve the cosmetic outcomes of LC.

The risk of conversion seems to be higher with SILC as compared to three and four port cholecystectomy. In a metaanalysis by Mate Milas et al, overall conversion of procedure was 69 (6%) among 1142 SILC [16]. The incidence of conversion with SILC was 4.39% vs. 0.53% with LC although the difference was statistically not significant (p value = 0.019). However with increasing experience with SILC the risk of procedure failure seems to have been reduced. In 10 trials with >40 SILC procedures, failure was 3.30% [17]. In our study there were 2 (10%) conversions in the SILC group as compared to 1 in conventional four port group(5%), and conversion rate was not significant (p=0.804). Conversion was mainly due to adhesions which interfered during dissection. Although the incidence appears to be higher than that reported in published literature but no definite inference could be drawn as the number of patients are quite less.

Milas M et al observed higher postoperative wound infection found following SILC. This may be due to longer periumbilical incision and its contamination during the delivery of the gallbladder, suboptimal hygiene of umbilicus itself despite cleaning [16,18]. Because anatomically umbilicus is probably the most difficult location for antiseptic and aseptic precautions and most SILC incision were given through umbilicus. Thus postoperative wound infection at the umbilical site has been a major concern [17] although infection seen was of minor SSI. In our study there has been a marginally higher incidence of wound infection but the difference was not statistically significant (p=0.765). Similar findings were also reported in metanalysis by Geng et al and Allemann et al [17,19]. During follow-up no incisional hernias were noted and it was ensured that sheath closure was done by the operating consultant. However, we need to have a longer follow-up to draw any inference on the development of incisional hernia from the present study.

We found in this study that the total operative time required for SILC (71.00 \pm 9.403) as compared to (39.50 \pm 9.162) in 4 port LC group which was significantly higher (p<0.001). This is in agreement with the metaanalysis by Liangyuan Geng et al which also concluded a longer operative time for SILC (p=0.005) [17]. Similar results were seen from other mataanalysis [20-23]. However the metaanlysis by Zhong et al including 7 RCT including 611 patients concluded that there was no significant difference in the operative time [24]. Similar metaanalysis by Lai EC et al [25] and Chang SK et al [26] did not find a significant difference in the operative time between the SILC and conventional LC groups.

Postoperative pain is a useful surrogate marker of procedure related trauma. It is often the predictor for early ambulation and return to work. SILC is being introduced as a less invasive procedure with lesser pain. Outcome of SILC in terms of postoperative pain is variable in literature. Meta-analysis of various studies suggest no difference in postoperative pain in both the techniques [16,17,19]. However studies included in these meta-analysis were often heterogenous and there was no uniformity in measure of pain. In our study we found no significant difference in the pain score at 8 hr after surgery (p=0.103), but on 7th day post operatively and after 3 months significantly low pain score was seen in SILC group. According to Geng et al there was no significant difference between post-operative pain in SILC and conventional laparoscopic cholecystectomy [17]. On the other hand, many studies showed a significant increase in post-operative [27-33]. On the other hand pain can be assessed by number of analgesics and nature of analgesics needed on the day of surgery. In our study patients undergone SILC require mostly single analgesics agent (p=0.223) and that too NSAIDS group (p=0.214). But these parameters are not statistically significant. As pain is felt differently for each patient, it is difficult to conclude on whether or not there is less post-operative pain in either of groups. In our study resuming daily work in SILC cases was significant (p=0.007) which may be due to decreased post operative pain, patients resumed in daily work early than the other two groups.

SILC was said to have a significantly shorter stay in the hospital [34-37]. Many though, didn't find a much significant difference in hospital stay [38-39]. In this study we did not find any significant difference in hospital stay among the two groups (p=0.223).

Cosmetic outcome is a very important parameter in assessing out come in laparoscopic surgeries. In a meta-analysis by Mate Milas et al, 5 trials with nonblinded patients (N=513) in favour of SILC (SMD=T.83, p value=0.037), but in 6 trials with blinded patients (N=719) difference was small and insignificant (SMD=0.42, p value=0.548) [16]. The reason for high cosmetic satisfaction score in SILC was attributed to the fact that the scars receded into the umbilicus and was hardly visible following SILC and patients were very satisfied with the cosmetic results. In our study, overall cosmetic satisfaction score was higher in SILC group. The cosmetic satisfaction was assessed at day 7(p<0.001), after 3 months (p<0.001) and 6 months (p<0.001).

In this study after 24 hr of surgery inflammatory parameter IL-6 found to be significantly lower in SILC group (p<0.001) than the 4 port LC group, which may explain the fast recovery in these cases by reducing surgical stress and infectious complications correlated to the surgical procedure. According to Luna et al serum IL-6 level after 6hr of surgery was found to lower in SILC group than conventional laparoscopic cholecystectomy group, but it was not statistically significant [39].

Conclusion

The SILC is associated with a longer operating time. This procedure has a lower incidence of early postoperative pain but no pain difference in immediate post-operative period. There is no additional complication associated with SILC as compared to the other 2 groups. There is no difference in the wound healing in three groups. Patients undergone SILC resume daily work early. SILC provide better cosmetic outcome. In SILC group postoperative inflammation is less which may explain the fast recovery in these cases by reducing surgical stress and infectious complications correlated to the surgical procedure. SILC is a safe and feasible procedure.

References

- 1. De U. Evolution of cholecystectomy: A tribute to Carl August Langenbuch. Indian J Surg 2004; 66: 97-100.
- 2. Mouret P. From the first laprospective futures. Dig Surg 1991; 8: 124.
- Santos BF, Teitelbaum EN, Arafat FO, Milad MP, Soper NJ, Hungness ES. Comparison of short-term outcomes between transvaginal hybrid NOTES cholecystectomy and laparoscopic cholecystectomy. SurgEndosc. 2012 Nov; 26(11): 3058-66.
- Auyang ED, Hungness ES, Vaziri K, Martin JA, Soper NJ. Human NOTES cholecystectomy: transgastric hybrid technique. J Gastrointest Surg. 2009; 13(6): 1149-50.
- Lehmann KS, Ritz JP, Wibmer A, et al. The German registry for natural orifice translumenal endoscopic surgery: report of the first 551 patients. Ann Surg 2010; 252: 263-70.
- Hodgett SE, Hernandez JM, Morton CA, Ross SB, Albrink M, Rosemurgy AS. Laparoendoscopic single site (Less) cholecystectomy. J Gastrointest Surg. 2009;

13(2): 188-92.

- Oruc MT, Ugurlu MU. Extraumbilical single-incision laparoscopic cholecystectomy with standard laparscopic instruments. Scand J Surg. 2013; 102(3): 209-14.
- Rao PP, Bhagwat SM, Rane A, Rao PP. The feasibility of single port laparoscopic cholecystectomy: a pilot study of 20 cases. HPB (Oxford). 2008; 10(5): 336-40.
- 9. Kravetz AJ, Iddings D, Basson MD, Kia MA. The learning curve with single-port cholecystectomy. JSLS. 2009; 13(3): 332-6.
- 10. Shussman N, Schlager A, Elazary R, Khalaileh A. Single-incisio Laparoscopic cholecystectomy: lessons learned for success. SurgEndosc 2010; 24(4): 1177-1179.
- Navarra G, Pozza E, Occhionorelli S, Carcoforo P, Donini I. One-wound laparoscopic cholecystectomy. Br J Surg 1997; 84(5): 695.
- Kulaylat MN, Dayton MT. Surgical Complications. In: Townsend CM, Beauchamp RD, Evers BM, Mattox KL, editors. Sabiston Textbook of Surgery, 18th ed. Philadelphia: Saunders: 2008; p. 331.
- 13. Zhu JF, Hu H, Ma YZ, Xu MZ, Li F. Transumbilical endoscopic surgery: a preliminary clinical report. SurgEndosc. 2009; 23(4): 813-7.
- 14. Strasberg SM, Hertl M, Soper NJ. An analysis of the problem of biliary injury during laproscopic cholecystectomy. J Am CollSurg 1995; 180: 101.
- 15. Thurley PD, Dhingsa R. Laparoscopic cholecystectomy: postoperative imaging. AJR Am J Roentgenol 2008; 191: 794.
- Milas M, Devidija S, Trkulja V. Single incision versus standard multiport laparoscopic cholecystectomy: Up-dated systematic review and meta-analysis of randomized trials. Surgeon. 2014 Feb 11; pii: S1479-666X(14)00015-18.
- 17. Geng L, Sun C, Bai J. Single incision versus conventional laparoscopic cholecystectomy outcomes: a meta-analysis of randomized controlled trials. PLoS One. 2013 Oct 2; 8(10): e76530.
- Sinan H, Demirbas S, Ozer MT, Sucullu I, Akyol M. Single-incision laparoscopic cholecystectomy versus laparoscopic cholecystectomy: a prospective randomized study. SurgLaparoscEndoscPercutan Tech 2012; 22(1): 12-6.
- Allemann P, Demartines N, Schafer M. Remains of the day: biliary complications related to single-port laparoscopic cholecystectomy. World J Gastroenterol. 2014 Jan 21; 203: 843-51.
- 20. Wang Z, Huang X, Zheng Q. Single-incision versus conventional laparoscopic cholecystectomy: a metaanalysis. ANZ J Surg. 2012 Dec; 82(12): 885-9.
- 21. Sajid MS, Ladwa N, Kalra L, Hutson KK, Singh KK, Sayegh M. Single-incision laparoscopic

cholecystectomy versus conventional laparoscopic cholecystectomy: metaanalysis and systematic review of randomized controlled trials. World J Surg 2012; 36(11): 2644-53.

- 22. Trastulli S, Cirocchi R, Desiderio J, Guarino S, Santoro A, Parisi A, Nova G, Boselli C. Systematic review and meta-analysis of randomized clinical trials comparing single-incision versus conventional laparoscopic cholecystectomy. Br J Surg 2013; 100: 191-208.
- 23. Arezzo A, Scozzari G, Famiglietti F, Passera R, Morino M. Is single-incision laparoscopic cholecystectomy safe? Results of a systematic review and meta-analysis. SurgEndosc. 2013 Jul; 27(7): 2293-304.
- 24. Zhong X, Rui YY, Zhou ZG. Laparoendoscopic singlesite versus traditional laparoscopic surgery in patients with cholecystectomy: a meta-analysis. J LaparoendoscAdvSurg Tech A. 2012 Jun; 22(5): 449-55.
- 25. Lai EC, Yang GP, Tang CN, Yih PC, Chan OC, Li MK. Prospective randomized comparative study of single incision laparoscopic cholecystectomy. Am J Surg. 2011 Sep; 202(3): 254-8.
- Chang SK, Wang YL, Shen L, Iyer SG, Shaik AB, Lomanto D. Interim report: a randomized controlled trial comparing postoperative pain in single-incision laparoscopic cholecystectomy and conventional laparoscopic cholecystectomy. Asian J Endosc Surg. 2013 Feb; 6(1): 14-20.
- Marks, J. Tacchino, R. Roberts, K. Onders, R. Denoto, G. Paraskeva, P.Shah. Prospective randomized controlled trial of traditional laparoscopic cholecystectomy versus single-incision laparoscopic cholecystectomy: report of preliminary data. The American Journal of Surgery, 2011; 3: 369-373.
- Lirici, M. M., Califano, A. D., Angelini, P., & Corcione, F. Laparo-endoscopic single site cholecystectomy versus standard laparoscopic cholecystectomy: results of a pilot randomized trial. The American Journal of Surgery, 2011; 202(1): 45-52.
- Ma J, Cassera MA, Spaun GO, Hammill CW, Hansen PD, Aliabadi-Wahle S. (2011). Randomized controlled trial comparing single-port laparoscopic cholecystectomy and four-port laparoscopic cholecystectomy. Annals of surgery, 2011; 254(1): 22-27.
- Phillips MS, Marks JM, Roberts K, Tacchino R, Onders R, DeNoto G, Shah S. Intermediate results of a prospective randomized controlled trial of

traditional four-port laparoscopic cholecystectomy versus single-incision laparoscopic cholecystectomy. Surgical endoscopy, 2012; 26(5), 1296-1303.

- Lee PC, Lo C, Lai PS, Chang JJ, Huang SJ, Lin MT, Lee PH. Randomized clinical trial of single incision laparoscopic cholecystectomy versus minilaparoscopic cholecystectomy. British Journal of Surgery, 2010; 97(7): 1007-1012.
- Ostlie DJ, Sharp NE, Thomas P, Sharp SW, Holcomb GW, St. Peter SD. Patient Scar Assessment After Single-Incision Versus Four-Port Laparoscopic Cholecystectomy: Long-Term Follow-Up from a Prospective Randomized Trial. Journal of Laparoendoscopic& Advanced Surgical Techniques, 2013; 23(6): 553-555.
- Moreira-Pinto J, Lima E, Correia-Pinto J, Rolanda C. Natural orifice transluminal endoscopy surgery: A review. World journal of gastroenterology: WJG, 2011; 17(33): 3795.
- 34. Antoniou SA, Pointner R, Granderath FA. Singleincision laparoscopic cholecystectomy: a systematic review. Surgical endoscopy, 2011; 25(2): 367-377.
- Gill ISA, Aron AP, Caddedu M, Canes J, Curcillo D, Teixeira PG. Consensus statement of the consortium for laparoendoscopic single-site surgery. Surgical endoscopy, 2010; 24(4): 762-768. http://dx.doi.org/ 10.1007/s00464-009-0688-8.
- Bucher P, Pugin F, Buchs NC, Ostermann S, Morel P. Randomized clinical trial of laparoendoscopic single site versus conventional laparoscopic cholecystectomy. British Journal of Surgery, 2011; 98(12): 1695-1702. http://dx.doi.org/10.1002/ bjs.7689.
- Gangl O, Hofer W, Tomaselli F, Sautner T, Függer R. Single incision laparoscopic cholecystectomy (SILC) versus laparoscopic cholecystectomy (LC)—a matched pair analysis. Langenbeck's Archives of Surgery, 2011; 396(6): 819-824.
- Barband A, Fakhree MB, Kakaei F, Daryani A. Singleincision laparoscopic cholecystectomy using glove port in comparison with standard laparoscopic cholecystectomy SILC using glove port. SurgLaparoscEndoscPercutan Tech. 2012; 22(1): 17-20.
- Luna RA, Nogueira DB, Varela PS, Rodrigues Neto Ede O, Norton MJ, RibeiroLdo C, Peixoto AM, de Mendonca YL, Bendet I, Fiorelli RA, Dolan JP. A prospective, randomized comparison of pain, inflammatory response, and short-term outcomes between single port and laparoscopic cholecystectomy. SurgEndosc. 2013 Apr; 27(4): 1254-9.