Impact of low- and standard-abdominal pressure on postoperative pain during laparoscopic cholecystectomy: A systematic review and meta-analysis of randomized trials

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Abstract

**Background:** This study aimed to comprehensively evaluate the impact of low- and standard-abdominal pressure on intraoperative, postoperative, and survival outcomes of Laparoscopic Cholecystectomy (LC).

**Methods:** A literature search of the databases, PubMed, Web of Science, Embase, and Cochrane Library was performed until April 30, 2021. Studies comparing low abdominal pressure and standard abdominal pressure for LC were included. Two reviewers independently screened the studies, extracted the data of interest, and assessed the risk of bias. Meta-analysis was performed using RevMan 5.3.

**Results:** Thirty-seven RCTs, including 2,104 patients, met the eligibility criteria. Low-abdominal pressure showed lower shoulder pain (RR: 0.55, 95% CI: 0.46, 0.67; p<0.001), lower analgesic use (RR: 0.54, 95% CI: 0.30, 0.99; p=0.039), lower pain score 0-6 h after operation (SMD: -0.28, 95% CI: -0.57, 0.00; p=0.04), lower pain score 7-12 h after operation (SMD: -0.81, 95% CI: -1.35, -0.28; p=0.003), lower pain score 13-24 h after operation (SMD: -0.66, 95% CI: -1.06, -0.26; p=0.001), longer operative time (WMD: 1.52, 95% CI: 0.26, 2.78, p =0.02), and length of hospital stay (WMD: -0.31, 95% CI: -0.54, -0.07; p=0.01). Conversion to open surgery (RR, 0.86; 95% CI: 0.28, 2.64; p=0.79) was not significantly different between the two groups.

**Conclusion:** Low abdominal pressure showed impressive improvement of shoulder pain and reduction of analgesic use. The pain score after surgery and length of hospital stay was also improved compared with standard abdominal pressure. The conclusions should be interpreted with caution because of the moderate to high degree of heterogeneity.
Introduction

Laparoscopic Cholecystectomy (LC) has been shown to be more beneficial than traditional cholecystectomy for patients with cholecystitis [1]. As a minimally invasive technique, LC improves the quality of postoperative living conditions and decreases the overall complications. LC is strongly recommended by the clinical guidelines worldwide [2]. However, some risk factors may affect the safety and efficacy of LC, especially abdominal pressure, which is considered to be one of the most important factors influencing postoperative complications [3].

For laparoscopic surgery, gas abdominal pressure caused by the input air is inevitable. Hence, postoperative complications, such as shoulder pain, may significantly affect the patient’s quality of life [4,5]. Moreover, postoperative pain is the most significant complication according to patients with LC [5]. To date, the relationship between abdominal pressure and pain is not clear. The severity of the postoperative pain is considered to be relative to the abdominal pressure [5].

In this review, abdominal pressure of LC was divided into two: Low Abdominal Pressure (LAP) and Standard Abdominal Pressure (SAP). Some minor corrections to the subgroups were made due to the information reported in the included studies [5]. According to recent LC guidelines, the pressure of the LAP and SAP groups was defined as 6–10 mmHg and 12–15 mmHg, respectively [5]. The latest meta-analyses [3,4] suggest that there is a lack of robust evidence on which type of abdominal pressure is beneficial for patients during LC.

Our study aimed to systematically evaluate and compare the impact of the two types of abdominal pressure on intraoperative, postoperative, and survival outcomes in patients who underwent laparoscopic cholecystectomy.

Materials and methods

Study registration

This study was registered on PROSPERO, and the registration number is CRD42018093851.

Study selection

We searched the databases, PubMed, Web of Science, Embase, and Cochrane Library, from their inception to May 31, 2019. The search was updated on April 30, 2021. Studies comparing low and standard abdominal pressure for LC were included. To search comprehensively and systematically for eligible studies, we also performed a manual search of the references found in the published reviews and articles.

Inclusion and exclusion criteria

The inclusion criteria for this meta-analysis were as follows: (1) randomized controlled trials (LAP versus SAP), (2) studies with a total sample size of more than 20, (3) studies where intraoperative and/or postoperative outcomes and/or survival outcomes were reported, and (4) LC patients. The exclusion criteria were as follows: (1) review articles, (2) correspondences or editorials, (3) conference abstracts without detailed data, (4) studies that included patients who underwent open cholecystectomy, (5) animal studies, and (6) single-arm studies.

Data extraction

Two authors (ZGR and LYF) independently reviewed all the identified studies. They resolved discrepancies by discussions. A third reviewer (ZZY) was consulted as necessary. We extracted the following items from each study: first author’s name, year of publication, country, publication type, and study type. There were no limitations in terms of language, race, and gender.

Outcomes of interest

The primary outcomes included shoulder pain, analgesic use, pain score 0-6 h after operation, pain score 7-12 h after operation, and pain score 13-24 h postoperatively. Secondary outcomes included conversion to open surgery, operative time, and length of hospital stay.

Statistical analysis

For dichotomous variables, Risk Ratios (RRs) were calculated with 95% Confidence Intervals (CIs). Continuous parameters, such as operative time and length of hospital stay, were analyzed using the Weighted Mean Difference (WMD). For outcomes such as pain scores, where different scales may have been used, the Standardized Mean Difference (SMD) was applied. We estimated the degree of heterogeneity among studies using the Cochrane Q statistic (p<0.10 was considered representative of statistically significant heterogeneity) and the I² statistic (I²>50% was considered to represent significant heterogeneity). Data were pooled using the random-effects model if significant heterogeneity among the studies was present. Otherwise, we used a fixed-effects model. We drew funnel plots to determine the possibility of publication bias when more than 10 studies were included. All data were analyzed using RevMan 5.3.

Results

Literature search

The search yielded 698 records. We excluded 65 studies because of duplication. We assessed the titles and abstracts of 633 studies and excluded 360 studies that did not meet the inclusion criteria. We assessed the full texts of 273 studies and fi-
nally included 37 studies for the present systematic review and meta-analysis. The flowchart is shown in Figure 1.

**Characteristics of eligible studies**

Thirty-seven RCTs and 2104 patients were included. All the included studies compared low- (1022 patients) to standard abdominal pressure (1082 patients) during LC. We redefined the LAP as 6-10 mmHg and SAP as 12-15 mmHg. Two RCTs presented data of patients from America, 11 RCTs provided data of patients from Europe, and 18 RCTs described patients from Asia. Of these, 17 lower abdominal pressure studies (≤8 mmHg) accounted for 50% of all the included studies. The analgesic procedures for all included studies were similar, in that general anesthesia was performed with fentanyl, remifentanil, or morphine. The details are presented in Table 1.

<table>
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<th>Study ID</th>
<th>Country</th>
<th>Study design</th>
<th>No. of patients</th>
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<th>Female (%)</th>
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LAP: Low-Abdominal Pressure; SAP: Standard-Abdominal Pressure; NA: Not Available; RCT: Randomized Controlled Trial.
Primary outcomes

Shoulder pain: Thirteen studies [10-12,19,20,23,24,27,29,32,34,39-42] including 1036 patients reported shoulder pain outcomes. Of the 505 patients in the LAP group, 104 patients suffered shoulder pain. In the SAP group of 527 patients, 201 reported shoulder pain. The meta-analysis showed that patients in the LAP group had lower shoulder pain than patients in the SAP group (RR: 0.55, 95% CI: 0.46, 0.67; p<0.00001). No statistical heterogeneity (I²=0.0%, p=0.520) was found (Figure 2).

Figure 2: Colonoscopy revealed mucosal diffuse congestion, oedema and scattered multiple irregular, deep chisel ulcers of the distal transverse colon, descending colon, sigmoid colon and rectum. The large mucosa of the sigmoid colon was absent and showed a mucosal island change. A: Descending colon; B and C: Sigmoid Colon; D: Rectum.

Analgesic use

Analgesic use was reported in nine studies [10,12,17,22,23,36,39] including 646 patients. Of the 326 patients in the LAP group, 100 used analgesics. In the SAP group of 320 patients, 162 used analgesics. The meta-analysis revealed that those in the LAP group had a lower analgesic use than patients in the SAP group (RR: 0.54, 95% CI: 0.30, 0.99; p=0.05). A moderate degree of heterogeneity (I²=89.0%, p<0.00001) was noted (Figure 3).

Figure 3: Comparison of analgesic use between LAP and SAP.

Pain score 0-6 h after operation

Nine studies [8,11,17,20,23,27,28,32] showed lower pain scores 0-6 h after operation in the LAP group (n=646, SMD: -0.28, 95% CI: -0.57, 0.00; p=0.05), with a moderate degree of heterogeneity (I²=65.0%, p=0.0001) (Figure 4).

Figure 4: Comparison of pain score 0-6 h after operation between LAP and SAP.

Secondary outcomes

Conversion to open surgery: Seven studies [8,9,21,23,28,35] with 315 patients revealed no statistical significance in the conversion to open surgery between the LAP and SAP groups (RR: 0.86, 95% CI: 0.28, 2.64; p=0.79), without statistical heterogeneity (I²=0.0%, p=0.85) (Figure 7).

Operative time: Thirty studies [6,8-20,22-28,32-35,37] with 1,777 patients reported the mean operative time. The LAP group had a longer operative time than the SAP group (WMD:1.52, 95% CI: 0.67, 2.38, p=0.001), with a low degree of heterogeneity (I²=26.0%, p=0.09) (Figure 8).

Figure 5: Comparison of pain score 7-12 h after operation between LAP and SAP.

Figure 6: Comparison of pain score 13-24 h after operation between LAP and SAP.

Figure 7: Comparison of the conversion to open surgery between LAP and SAP.

Figure 8: Comparison of operative time between LAP and SAP.

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Length of hospital stay: The length of hospital stay was reported in eight studies [7,8,10,11,22,24,32] (655 patients). The LAP group showed a shorter length of hospital stay than the SAP group (WMD: -0.31, 95% CI: -0.54, -0.07; p<0.00001) (Figure 9).

Operative time and length of hospital stay were compared between the two groups. Patients in the LAP group had a shorter operative time and length of hospital stay. The length of hospital stay was presented with a high degree of heterogeneity as most studies that did not define the length of hospital accurately and concisely. Moreover, the two groups had no significant difference in the conversion to open surgery, which showed no benefit in either group.

Surprisingly, shoulder pain, pain score 0-6 h after operation, pain score 7-12 h after operation, pain score 13-24 h after operation were associated with a high degree of heterogeneity due to some inevitable bias; hence, the results should be interpreted with caution.

Figure 8: Comparison of operative time between LAP and SAP.

Figure 9: Comparison of the length of hospital stay between LAP and SAP.

Publication bias

Three outcomes (shoulder pain, pain score 13-24 h after operation, and operative time) included more than 10 studies. Funnel plots suggested that the possibility of publication bias was small.

Discussion

This meta-analysis indicated that the LAP group was more beneficial for lower shoulder pain, analgesic use, pain score 0-6 h after operation, pain score 7-12 h after operation, pain score 13-24 h after operation, and length of hospital stay. Although the operative time of the LAP group was longer. Other outcomes, such as conversion to open surgery, were not significantly different between the two types of abdominal pressure for LC. Most of the results attained a moderate degree of heterogeneity due to some inevitable bias; hence, the evidence may not be robust enough for clinical practice.

Interestingly, the LAP group had lower shoulder pain than the SAP group in most of the included studies, which suggests that there was no heterogeneity. Shoulder pain results from the pressure elicited by the injected gas in the abdomen. Although LC is widely popular worldwide, the possible postoperative complications remain confusing to many clinical workers [43]. Shoulder pain is one of the inevitable postoperative complications that influence the patient’s quality of life in a large part. Hence, it is necessary to choose a favorable abdominal pressure to reduce the incidence of shoulder pain. Given these results, LAP may minimize shoulder pain.

The LAP group also indicated a lower pain score 0-6 h after the operation, a pain score of 7-12 h, and a pain score of 13-24 h postoperatively. Pain score was evaluated using the visual analog scale (VSA) score, where 0 represents no pain and 10 represents maximum pain. VSA assesses the severity of post-operative complications approximately [44]. A high degree of heterogeneity may be generated due to different ways of evaluating pain in the included studies. The outcomes of pain score 0-6 h, pain score 7-12 h after operation, and pain score 13-24 h after operation were associated with a high degree of heterogeneity. Hence, the results should be interpreted with caution.

Analgesic use was also evaluated in this meta-analysis. The LAP group used less analgesic than the SAP group. This was associated with a moderate degree of heterogeneity. Analgesics are commonly used to relieve the discomfort caused by pain. Although LC is minimally invasive, postoperative pain does occur to some extent. The lower incidence of analgesic use may improve the quality of life for the patients with LC.

Operative time and length of hospital stay were compared between the two groups. Patients in the LAP group had a shorter operative time and length of hospital stay. The length of hospital stay was presented with a high degree of heterogeneity as most studies that did not define the length of hospital accurately and concisely. Moreover, the two groups had no significant difference in the conversion to open surgery, which showed no benefit in either group.

Recently published meta-analysis analyses [3,4] reported that LAP is more beneficial than SAP but the outcomes were not analyzed systematically. We included the newly published RCTs from 2014 to 2018 and analyzed the data comprehensively in this meta-analysis. The results indicated that LAP is not inferior to SAP. Furthermore, LAP may lead to better safety and efficacy. Well-conducted and updated meta-analyses of RCTs are accepted as the best-quality evidence for informing clinical practice and health policy [45,46]. However, there are still some unsolved questions when comparing LC using LAP versus SAP. The incidence of malignant heart events was not shown in this meta-analysis. The operative time and length of hospital stay were also shortened in the LAP group, suggesting that LAP may generate fewer complications and enhance the recovery of patients after LC.
The findings of this study indicated that LAP has obvious advantages compared with SAP, although most outcomes were associated with a high degree of heterogeneity. The heterogeneity may come from different pain evaluation methods, different populations, different operations performed by different surgeons, and so on. In sum, LAP for LC improves intraoperative and postoperative complications, without affecting the surgeons’ visual and physical access. In other words, lower pressure may provide a wide and clear view during the operation.

Considering the recent clinical guidelines [48,49] on laparoscopic surgery, a few have recommended LAP for patients with gall bladder diseases. Previous studies [6-8] have suggested that LAP may not provide an obvious and high-quality view of the operative field because low abdominal pressure may not provide enough space for the surgeons. In contrast, according to our results, the use of LAP for the operation did not hinder access, regardless of the heterogeneity, and that LAP may not offer a larger operative space. To the best of our knowledge, LAP provides adequate space for the surgeons, as well as better safety and efficacy. However, our results need to be validated by high-quality prospective studies.

There are some limitations in this meta-analysis. (1) Some heterogeneity may influence the results to some extent. We did not perform the certainty of evidence assessment using the GRADE system because of the lack of skill and experience with the GRADE method [50]. (2) Some of the included studies had inevitable biases, which may affect the quality of the evidence. (3) Due to sufficient data, some outcomes could not be meta-analyzed to offer more comprehensive evidence for LC.

Some merits were also shown in this meta-analysis. (1) All the included studies were randomized controlled trials comparing LAP and SAP. (2) Almost all the necessary outcomes, including intraoperative and postoperative complications, were analyzed in this meta-analysis. (3) This is the first meta-analysis comparing LAP and SAP with systematic data and comprehensive RCTs published until 2019. In addition, our search was guided by an experienced information expert who improved the search quality [51].

Conclusions

LAP showed impressive improvement of shoulder pain and reduction of analgesic use. LAP was also associated with improved pain score after surgery and length of hospital stay compared with that of standard abdominal pressure. The conclusions should be interpreted with caution, and in combination with clinical practice experience, because most of the results had a high degree of heterogeneity.

Declarations

Conflict of interest: The authors have no conflicts of interest to declare.

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