

Systematic Review Comparing Early Versus Delayed Cholecystectomy for Acute Cholecystitis

¹Ahmad Adnan Hariri, ²Hatoon Mohammed Alkholi, ³Rasha Saeed Taher, ⁴Tahr Abdullah Alqurashi, ⁵Abdullah Ahmed Alwayrqi, ⁶Ahmed Maroof Ba Dheeb, ⁷Mohammad Abduh Algozi

Abstract: Acute cholecystitis is one of the most substantial diseases among population, takes place most frequently due to blockage of the cystic duct with gallstones (cholelithiasis) and is among the most common severe abdomen diseases in emergency clinic. This systematic review was aimed to compare early cholecystectomy for acute cholecystitis versus delayed cholecystectomy, we also intended to overview the benefits and the disadvantages in between two procedures. The MIDLINE, Embase, Cochrane Library and Central Register of Controlled Trials (CENTRAL) databases were searched from inception to December 2016. The electronic searches were performed using exploded medical Mesh terms including ‘cholecystitis’, ‘cholecystectomy’ and” acute cholecystitis” “early” and “delayed”. The searches were limited to human subjects and English language restriction was applied. The references of relevant reviews and included studies were also searched for more potentially eligible studies to our concerned topic. Our findings revealed no risk distinction between early and delayed surgical treatment on the basis of results in mortality, morbidity, and rates of conversion. The mean total health center stay was much shorter in the early group than in the postponed group, and there was no difference in operation time in between the two groups. cholecystectomy within 24 hours of healthcare facility admission was revealed to be superior to the conservative method worrying morbidity.

Keywords: Gallstones (cholelithiasis), Controlled Trials (CENTRAL), MIDLINE, Embase.

1. INTRODUCTION

Acute cholecystitis is one of the most substantial diseases among population, takes place most frequently due to blockage of the cystic duct with gallstones (cholelithiasis) and is among the most common severe abdomen diseases in emergency clinic. About 5 - 25% of the adult Western population have gallstones, patients with gallstones and older adults are affected ⁽¹⁾. Since the risk of establishing subsequent episodes of cholecystitis is high, laparoscopic cholecystectomy is typically suggested for acute cholecystitis. Controversy exists about the finest timing for surgery ⁽²⁾. Generally, two approaches are pursued: early surgery versus an initial conservative treatment with prescription antibiotics for total resolution of inflammation, followed by postponed laparoscopic cholecystectomy numerous weeks later ⁽²⁾. Delayed surgery is based on the presumption that affected inflammatory tissue is more susceptible to surgical interventions and results in an increased risk of surgical complications. Throughout its early years, laparoscopic cholecystectomy was contraindicated in intense cholecystitis ⁽³⁾. In support of the postponed approach, a single-center cost-utility analysis favored standard management of acute cholecystitis over early cholecystectomy since of lower incremental expense per quality-adjusted life year got (4). More current research studies do not support the necessity of the conservative pretreatment to conquer the intense inflammatory response ^(5,14). The waiting period might be associated with greater morbidity ⁽⁷⁾. The ideal timing of laparoscopic cholecystectomy for severe cholecystitis is arguable ^(8,9). Previous systemic evaluations ^(10,11,12) have revealed that early laparoscopic cholecystectomy (ELC) is as safe as postponed laparoscopic cholecystectomy (DLC). ELC for acute cholecystitis is associated with a shorter health center stay. However, these

incomplete analyses typically paid little focus on health center expenses, work days lost and quality of life, which may puzzle and underestimate the differences^(10,11,12).

This systematic review was aimed to compare early cholecystectomy for acute cholecystitis versus delayed cholecystectomy, we also intended to overview the benefits and the disadvantages in between two procedures.

2. METHODOLOGY

The study (*systematic review*) was conducted according to the recommendations of the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines for reporting meta-analyses⁽¹³⁾.

Search strategy:

The MIDLINE, Embase, Cochrane Library and Central Register of Controlled Trials (CENTRAL) databases were searched from inception to December 2016. The electronic searches were performed using exploded medical Mesh terms including 'cholecystitis', 'cholecystectomy' and "acute cholecystitis" "early" and "delayed". The searches were limited to human subjects and English language restriction was applied. The references of relevant reviews and included studies were also searched for more potentially eligible studies to our concerned topic.

Authors executed the literature search independently, removed the duplicate records, screened the titles and abstracts for relevance, and tagged the articles as included, excluded or requiring further assessment.

3. RESULTS AND DISCUSSION

Operating time between the two groups (early & delayed):

There was no significant distinction in the operating time between the two groups (MD -1.22 minutes; 95% CI -3.07 to 0.64). There was significant heterogeneity ($I^2 = 76\%$; $P = 0.001$). Embracing the random-effects model showed that the operating time was considerably longer in the early group than the postponed group (MD 15.31 minutes; 95% CI 1.09 to 29.53). Two trials reported the mean^(14,15), one trial reported the mean and standard deviation of the operating times in the people who went through successful laparoscopic cholecystectomy and in those who went through conversion to open surgery separately⁽¹⁶⁾, and three trials reported the mean operating time^(17,18,19). The average was used in the meta-analysis. The median operating times reported in the two of the three trials was 21 minutes and 30 minutes longer in the early group than the delayed group^(17,18). The mean operating time in one trial in which laparoscopic common bile duct exploration was used for thought common bile duct stones on routine peri-operative cholangiogram (with surgical locals carrying out these treatments) was 2 minutes much shorter in the early group⁽¹⁹⁾, in this trial, there was no substantial heterogeneity noted ($I^2 = 0\%$; $P = 0.76$), and the total operating time was substantially longer in the early group than in the delayed group (MD 20.06 minutes; 95% CI 10.21 to 29.90).

There was considerable heterogeneity amongst these trials with others. In the other tracks There was a considerable difference in operation time between the two groups (MD 15.31 (95% CI 1.09 to 29.53). 2 trials reported the mean^(14,17) and 3 trials reported the mean operating time^(15,18,19).

Total Hospital Stay between the two groups:

Five research studies reported this result^(14,15,16,17,18). The mean and average were used in different research studies. The typical was used in the meta-analysis after assigning the standard deviation from the worth^(15,17). Overall medical facility stay was significantly decreased in ELC group versus DLC group (MD - 4.12 days (95% CI, - 5.22 to - 3.03). There was no heterogeneity (%). The outcome remained if we excluded two median-using trials. In Gutt et al.'s trial⁽⁶⁾, imply with interquartile was utilized and could not be applied to meta-analysis. Mean total length of health center stay for immediate LC group was 5.4 days versus 10.0 days for DLC group. Length of healthcare facility stay after cholecystectomy was about the same in both groups. Thus, there might be no significant distinction in surgical complications for various timing.

Bile duct injury and other complications:

Five trials reported bile duct injury needing re-operations^(15,16,17,18,19). There was no substantial distinction between the two groups for this most feared complication (Peto OR 0.49; 95% CI 0.05 to 4.72). There was no substantial heterogeneity ($I^2 = 28\%$; $P = 0.25$). There was no change in the interpretation of outcomes using the random-effects model or the risk difference. Imputing the results for the one trial where there were post-randomisation drop-outs did not

change the interpretation of the findings in three of the 4 circumstances. In the best-worst scenario, the percentage of individuals who developed bile duct injury was substantially lower in the early group than the delayed group (Peto OR 0.17; 95% CI 0.05 to 0.54). There was no considerable difference between the groups in the other situations. Other Complications were stated in Gutt et al.'s study⁽⁶⁾ provided that morbidity in ELC group was substantially lower than in DLC group (11.8% versus 34.4%). There was no substantial difference in between the 2 groups in basic (RR 0.72 (95% CI 0.36 to 1.46). There was heterogeneity amongst this trial with others.

Gallstone-Related Morbidity during Waiting Time. During waiting periods of DLC group, 33 patients developed cholangitis and 3 got pancreatitis (cholangitis: 31/314; pancreatitis: 3/314, in Gutt et al.'s research study⁽⁶⁾. In five of the included studies^(15,16,17,18,19), 17.5% (40/228) of patients in DLC group had actually repeated signs or were not relieved throughout the waiting period^(14,15,17). Conversion to Open Cholecystectomy(OC). All the research studies consisted of conversion to OC. There was no considerable difference in conversion to OC in between the two groups (RR 0.91 (95% CI 0.69 to 1.20). The conversion rate was 14.4% (79/548) in ELC and 15.6% (87/558) in DLC group. There was no considerable heterogeneity (%).

Patient satisfaction and quality of life:

The total rate of satisfaction was higher for patients in the early groups than for those in the delayed group⁽¹⁹⁾. Johansson et al⁽¹⁸⁾ reported ELC to be connected with a more favorable intestinal sign rating at 1 month after surgical treatment, whereas there were no significant distinctions at 3 and 6 months. Macafee et al⁽⁴⁾ reported considerably lower visual analogue scale ratings in the ELC group compared to the DLC group, however both treatments led to rapid healing. On the other hand, Bile duct injury is the most feared complication throughout cholecystectomy and can be deadly⁽²⁰⁾. Restorative surgical treatment for bile duct injury has a high morbidity and death^(20,21) and the lifestyle can be bad, even three years after corrective surgical treatment⁽²²⁾. Cholecystitis has actually been thought about as a risk factor for bile duct injury⁽²¹⁾. Observational studies have suggested a greater variety of bile duct injuries with early surgery^(17,18). This difference could be due to the absence of 'intention-to-treat' analysis in observational studies, with individuals from the delayed surgery group prepared to go through emergency surgical treatment being consisted of in the early surgical treatment group, that is, the performance of an unsuitable 'treatment-received analysis'. Larger research studies are required to demonstrate the little differences in bile duct injury rates between an early or delayed approach to acute cholecystitis. Nevertheless, it is unlikely that large trials will be performed as the needed sample size that would allow to assess differences in the bile duct is high; the 5 trials offering data for this meta-analysis contributed to much less than 1% of the individuals' needed sample size. There were no considerable differences in the other major issues related to surgical treatment between the two groups.

The optimal timing of surgical treatment was still controversial. The evaluated trials do not have a homogeneous definition of early surgical treatment. "Early" has actually been variably specified as anywhere from 24 h to 7 days after either the start of signs or the time of medical diagnosis at health center admission according to the consisted of trials. Lee et al⁽²³⁾ suggested the ideal threshold period must be within 72 h of symptom start. In our evaluation, a subgroup analysis revealed no considerable difference in the proportion of people with conversion to OC or problems in between people operated on within 4 days and within 7 days after the beginning of symptoms. More data concerning the optimal timing is needed for further analysis.

The total health center stay of the individuals was much shorter by 4 days with ELC than with postponed surgery. This might arise from the more treatments and treatments. Surgical treatment can clear the inflammation tissue and abscess much quicker than conservative therapy. Another reason was that patients in delayed group might receive extra treatment when the syndromes were not relieved or repeated. Gurusamy et al.⁽²⁴⁾ also discussed that lack of blinding could be an essential source of predisposition in the length of hospital stay.

4. CONCLUSION

Our findings revealed no risk distinction between early and delayed surgical treatment on the basis of results in mortality, morbidity, and rates of conversion. The mean total health center stay was much shorter in the early group than in the postponed group, and there was no difference in operation time in between the two groups. cholecystectomy within 24 hours of healthcare facility admission was revealed to be superior to the conservative method worrying morbidity. We believe that immediate cholecystectomy needs to end up being treatment of option for acute cholecystitis in operable patients. In contrast there was no considerable distinction between early and late laparoscopic cholecystectomy on our

main results. trials with high risk of bias show that early cholecystectomy throughout acute cholecystitis seems safe and might reduce the total hospital stay.

REFERENCES

- [1] Halldestam, E.-. Enell, E. Kullman, and K. Borch, "Development of symptoms and complications in individuals with asymptomatic gallstones," *British Journal of Surgery*, vol. 91, no. 6, pp. 734–738, 2004.
- [2] Papi C, Catarci M, D'Ambrosio L, et al. Timing of cholecystectomy for acute calculous cholecystitis: a meta-analysis. *Am J Gastroenterol*. 2004;99:147– 155.
- [3] Wilson P, Leese T, Morgan WP, et al. Elective laparoscopic cholecystectomy for "all-comers." *Lancet*. 1991;338:795–797.
- [4] Macafee DA, Humes DJ, Bouliotis G, et al. Prospective randomized trial using cost-utility analysis of early versus delayed laparoscopic cholecystectomy for acute gallbladder disease. *Br J Surg*. 2009;96:1031–1040.
- [5] Johansson M, Thune A, Blomqvist A, et al. Management of acute cholecystitis in the laparoscopic era: results of a prospective, randomized clinical trial. *J Gastrointest Surg*. 2003;7:642–645.
- [6] C. N. Gutt, J. Encke, J. Königer, et al., "Acute cholecystitis: early versus delayed cholecystectomy, a multicenter randomized trial (ACDC study, NCT00447304)," *Annals of Surgery*, vol. 258, no. 3, pp. 385–393, 2013. Germanos S, Gourgiotis S, Kocher HM. Clinical update: early surgery for acute cholecystitis. *Lancet*. 2007;369:1774–1776.
- [7] Yamashita Y, Takada T, Strasberg SM, Pitt HA, Gouma DJ, Garden OJ et al. TG13 surgical management of acute cholecystitis. *J Hepatobiliary Pancreat Sci* 2013; 20: 89–96.
- [8] Yamashita Y, Takada T, Hirata K. A survey of the timing and approach to the surgical management of patients with acute cholecystitis in Japanese hospitals. *J Hepatobiliary Pancreat Surg* 2006; 13: 409–415.
- [9] Shikata S, Noguchi Y, Fukui T. Early versus delayed cholecystectomy for acute cholecystitis: a meta-analysis of randomized controlled trials. *Surg Today* 2005; 35: 553–560.
- [10] Lau H, Lo CY, Patil NG, Yuen WK. Early versus delayed-interval laparoscopic cholecystectomy for acute cholecystitis: a meta-analysis. *Surg Endosc* 2006; 20: 82–87.
- [11] Siddiqui T, MacDonald A, Chong PS, Jenkins JT. Early versus delayed laparoscopic cholecystectomy for acute cholecystitis: a meta-analysis of randomized clinical trials. *Am J Surg* 2008; 195: 40–47.
- [12] Moher D, Liberati A, Tetzlaff J, Altman DG; PRISMA Group. Preferred reporting items for systematic reviews and meta-analyses: the PRISMA statement. *Int J Surg* 2010; 8: 336–341.
- [13] Kolla SB, Aggarwal S, Kumar A, Kumar R, Chumber S, Parshad R, et al. Early vs delayed laparoscopic cholecystectomy for acute cholecystitis. *Surgical Endoscopy* 2004;18:1323–7.
- [14] Lai PBS, Kwong KH, Leung KL. Randomized trial of early versus delayed laparoscopic cholecystectomy for acute cholecystitis. *British Journal of Surgery* 1998;85(6):764–7.
- [15] Yadav RP, Adhikary S, Agrawal CS, Bhattarai B, Gupta RK, Ghimire A. A comparative study of early vs. delayed laparoscopic cholecystectomy in acute cholecystitis. *Kathmandu University Medical Journal* 2009;7(25):16–20.
- [16] Lo C, Liu C, Fan S, Lai ECS, Wong J. Prospective randomized study of early versus delayed laparoscopic cholecystectomy for acute cholecystitis. *Annals of Surgery* 1998;227(4):461–7.
- [17] Davila D, Manzanares C, Picho ML, Albors P, Cardenas F, Fuster E, et al. Experience in the treatment (early vs. delayed) of acute cholecystitis via laparoscopy. *Cirugia Espanola* 1999;66(Suppl 1):233.
- [18] Johansson M, Thune A, Blomqvist A, Nelvin L, Lundell L. Impact of choice of therapeutic strategy for acute cholecystitis on patient's health-related quality of life: results of a randomized, controlled clinical trial. *Digestive Surgery* 2004;21(5-6):359–62.
- [19] Saber A, Hokkam EN. Operative outcome and patient satisfaction in early and delayed laparoscopic cholecystectomy for acute cholecystitis. *Minim Invasive Surg* 2014; 2014: 162643.

- [20] Sicklick JK, Camp MS, Lillemoe KD, Melton GB, Yeo CJ, Campbell KA, et al. Surgical management of bile duct injuries sustained during laparoscopic cholecystectomy: perioperative results in 200 patients. *Annals of Surgery* 2005; 241(5):786–95.
- [21] Schmidt SC, Langrehr JM, Hintze RE, Neuhaus P. Longterm results and risk factors influencing outcome of major bile duct injuries following cholecystectomy. *British Journal of Surgery* 2005;92(1):76–82.
- [22] Moore DE, Feurer ID, Holzman MD, Wudel LJ, Strickland C, Gorden DL, et al. Long-term detrimental effect of bile duct injury on health-related quality of life. *Archives of Surgery* 2004;139(5):476–82
- [23] A. Y. Lee, J. J. Carter, M. S. Hochberg, A. M. Stone, S. L. Cohen, and H. L. Pachter, “The timing of surgery for cholecystitis: a review of 202 consecutive patients at a large municipal hospital,” *The American Journal of Surgery*, vol. 195, no. 4, pp. 467–470, 2008.
- [24] K. S. Gurusamy, C. Davidson, C. Gluud, and B. R. Davidson, “Early versus delayed laparoscopic cholecystectomy for people with acute cholecystitis,” *Cochrane Database of Systematic Reviews*, vol. 6, Article ID CD005440, 2013.