

Factors Associated with Moderate to Severe Pain after Laparoscopic Surgery

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Abstract:

Objective: The aim of this study was to describe associated factors of patients with moderate to severe pain after laparoscopic abdominal surgery in Post Anesthetic Care Unit.

Material and Methods: This retrospective study included patients who underwent laparoscopic abdominal surgeries (bariatric, colorectal, gynecological, urological, and upper gastrointestinal surgery) between February and July 2019. Demographic data were retrieved from the anesthetic records. Pain score was evaluated by self-rating using verbal numerical rating scale (0–10). Logistic regression was used to obtain independent risk factors of moderate to severe (score 4–10) pain against no or mild pain.

Results: Two-hundred and ten patients were included. The incidence of moderate (score 4–6) to severe pain (score 7–10) was 50.0%. Those who suffered from moderate to severe pain required higher consumption of analgesic drugs in recovery room than the other groups, however durations of hospitalization were not different between the two groups (p -value=0.329). This was because the highest intensity of pain was in the first postoperative day. A multivariate logistic regression analysis revealed that abdominal pressure more than 12 mmHg (odd ratio (OR)=1.84, 95% confidence interval (CI) 0.8–4.22, p -value=0.153), and operation time more than 3 hours (OR 1.49, 95% CI 0.86–2.61, p -value=0.158) were independent risk factors of moderate to severe pain after laparoscopic procedure

Conclusion: High intra-abdominal pressure (≥ 12 mmHg) and prolonged operation time (>3 hours) are the independent risk factors of moderate and severe pain in laparoscopic surgery.

Keywords: factors, moderate to severe pain, laparoscopic surgery

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Introduction

Postoperative pain is a common post-surgical problem, which prolongs recovery and delays hospital discharge.¹ Consequently, surgeons have been persuaded to apply surgical techniques with minimal invasiveness in order to reduce postoperative pain, one of which is laparoscopic surgery. However, moderate to severe pain are also reported after laparoscopic surgical procedures, and a previous study reported that laparoscopic surgery had higher analgesic requirement than laparotomy surgery.² The frequency of moderate to severe pain has been demonstrated as high as 50.0–60.0%.^{3,4} Therefore the knowledge regarding factors associated moderate to severe pain in laparoscopic surgeries is important for controlling pain.

Postoperative pain is likely related to multifactorial factors including patient factors, surgical factors, and anesthetic factors. Previous literature has reported and focused only on patient factors. These studies reported young age⁴, pre-existing pain⁵, female^{6,7}, and obese patients which are patient factors associated pain. However few studies documented surgical factors including duration of surgery, port site and intra-abdominal pressure^{8,9}, which would be correlated with pain after laparoscopic procedures. Moreover, anesthetic factors such as local infiltration anesthesia¹⁰, paracetamol¹¹, and parecoxib¹² would be the prophylactic factors to reduce pain.

Regarding the aforementioned information, there is a lack of information about surgical and anesthetic factors in abdominal surgery. To fulfil this knowledge gap, the aim of this study was to describe associated factors and outcomes of patients with moderate to severe pain after any type of laparoscopic surgery.

Material and Methods

This was a retrospective cross-sectional study. The research retrieved the records of 210 patients which underwent laparoscopic surgery (bariatric, colorectal (laparoscopic low anterior resection), gynecological, urological, and upper GI surgery (laparoscopic cholecystectomy, laparoscopic gastrectomy) in Songklanagarind Hospital, a tertiary hospital in southern Thailand between February and July 2019. The inclusion criteria were age ≥ 18 years old, laparoscopic surgery, and American Society of Anesthesiologists (ASA) classification I–III. Exclusion criteria were converting to explore laparotomy, and emergency case. All data (demographic data, ASA classification, type of surgery, duration of surgery, and pain score at recovery room) were extracted from anesthetic records in the Anesthesiology department. Consequently, we reviewed and obtained port site operation, abdominal pressure from operative notes, duration of hospitalization, and opioid consumption in the 24 hour post-operative period from electronic medical records of this hospital. The primary outcome of this study was pain evaluated by the patient based on verbal numerical rating scale (0–10) at Post Anesthetic Care Unit after laparoscopic surgery. The scores were categorized into no pain (0), mild pain (1–3), moderate pain (4–6), and severe pain (7–10). Then patients were divided into two groups, no/mild pain and moderate to severe pain. The secondary outcomes were opioid consumption in the recovery room, and 24 hour postoperative period, and duration of hospitalization among groups.

The sample size was calculated based on the difference in a previously reported proportion of moderate/severe pain in obesity patients [proportion (p)=0.43] and the proportion of mild pain in obesity patients (p=0.23).⁷ Using a two-sided test with a 95% confidence interval (CI)

and type II error of 0.2, the calculated sample size was 96 patients/group. The number was increased by 10.0% drop out to compensate for excluded cases.

Statistical analysis was performed using program R. Categorical variables such as gender, body mass index (BMI), ASA classification, type of laparoscopic surgery, type of intraoperative opioid, paracetamol and parecoxib use were demonstrated as percentage and proportion. Continuous variables such as weight, height, duration of hospitalization, and opioid consumption were presented as mean and standard deviation or median and interquartile range. The parameters were compared between moderate/severe pain and no/mild pain group by Student t-test

or Wilcoxon Rank Sum test as appropriate. Categorical variables were analyzed using chi-squared test or Fisher's Exact test as appropriate. Variables with p -value < 0.200 were included in multivariate logistic regression model. A p -value less than 0.050 was considered statistically significant.

Results

A total of 212 patients were assessed for study eligibility, 210 patients were retrieved in this study. Two patients were excluded from this study due to converting to explore laparotomy. Hence 105 patients of each group were available for analysis (Figure 1).

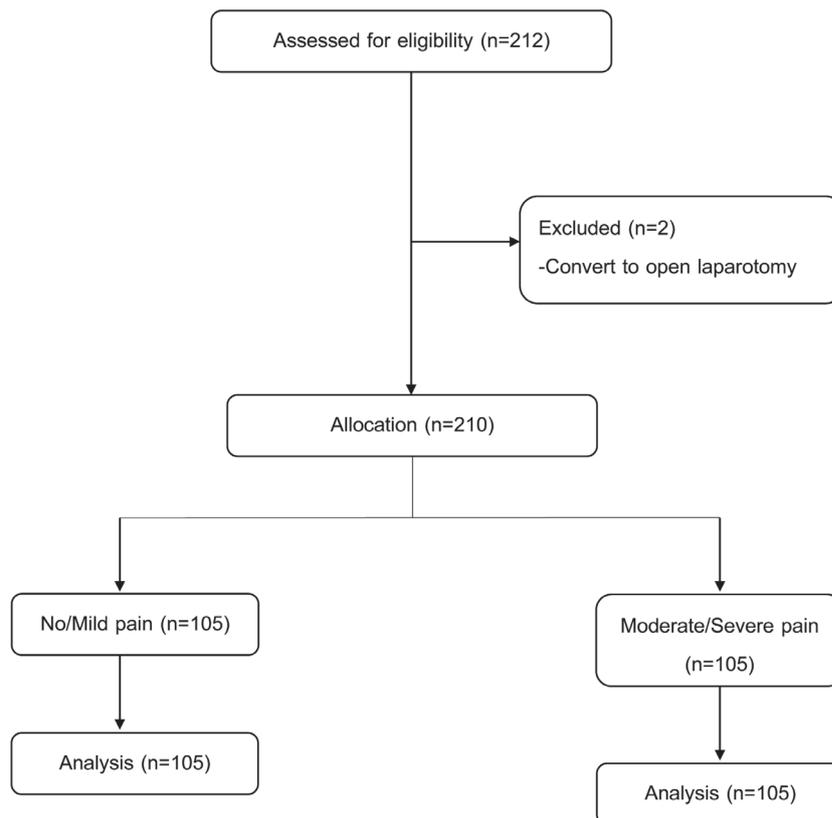


Figure 1 Consort flow diagram

Table 1 Characteristic data of patients in laparoscopic surgery

Variables	No/mild pain (n=105)	Moderate/severe pain (n=105)	p-value
Age (years)			
≤40	32 (47.8)	35 (52.2)	0.767
>40	73 (51.0)	70 (49.0)	
Weight (kg), median (IQR)	60 (51.0, 70.0)	65 (57.0, 87.0)	0.005
Height (cm), mean (S.D.)	159.3 (8.4)	161.3 (8.0)	0.076
BMI (kg/m ²)			
<35	92 (53.8)	79 (46.2)	0.033
≥35	13 (33.3)	26 (66.7)	
Sex			
Male	35 (51.5)	33 (48.5)	0.883
Female	70 (49.3)	72 (50.7)	
ASA classification			
I	8 (57.1)	6 (42.9)	0.679
II	70 (51.1)	67 (48.9)	
III	27 (45.8)	32 (54.2)	
Technic of anesthesia			
GA alone	104 (49.8)	105 (50.2)	1.000
GA combine RA	1 (100.0)	0 (0.0)	
Type of surgery			
Bariatric surgery	9 (29.0)	22 (71.0)	0.124
Gynecological surgery	38 (55.1)	31 (44.9)	
Upper GI surgery	28 (54.9)	23 (45.1)	
Colorectal surgery	26 (53.1)	23 (46.9)	
Urological surgery	4 (40.0)	6 (60.0)	
Abdominal pressure (mmHg)			
Pressure<12 mmHg (P mean=10 mmHg)	17 (63.0)	10 (37.0)	0.216
Pressure≥12 mmHg (P mean=15 mmHg)	88 (48.1)	95 (51.9)	
Duration of surgery (mins)			
≤180	69 (53.9)	59 (46.1)	0.203
>180	36 (43.9)	46 (56.1)	
Port			
Single port	8 (53.3)	7 (46.7)	1.000
Multiple ports	97 (49.7)	98 (50.3)	
Local infiltration			
Yes	97 (50.8)	94 (49.2)	0.630
No	8 (42.1)	11 (57.9)	
Types Opioid intraoperative use			
Fentanyl	80 (49.7)	81 (50.3)	0.062
Morphine	25 (51.0)	24 (49.0)	
Paracetamol IV intraoperative			
Yes	14 (48.3)	15 (51.7)	1.000
No	91 (50.3)	90 (49.7)	
NSAIDs IV intraoperative			
Yes	23 (46.0)	27 (54.0)	0.627
No	82 (51.2)	78 (48.8)	

Data are presented as number (%), unless indicated otherwise.

kg=kilogram, IQR=interquartile range, cm=centimeter, S.D.=standard deviation, BMI=body mass index, ASA=American Society of Anesthesiologists, GA=general anesthesia, RA=regional anesthesia, GI=gastrointestinal, P=pressure, mm=millimeter, Hg=mercury, min=minute, NSAIDs=non-steroidal anti-inflammatory drugs, IV=intravenous

Table 1 demonstrated no statistically significant differences in demographic data such as age, sex, ASA physical status, technique of anesthesia, and type of laparoscopic surgery among groups. Nonetheless moderate to severe pain group had a number of obese patients ($BMI \geq 35$), more significantly than other groups (average $BMI = 25 \text{ kg/m}^2$, and 23.7 kg/m^2 , respectively ($p\text{-value} = 0.036$)).

Table 2 displayed fentanyl, and morphine consumption in recovery room, and morphine consumption in 24 hours were found higher in moderate to severe pain group than other groups, $p\text{-value} < 0.001$. In addition, paracetamol and non-steroidal anti-inflammatory drugs (NSAIDs) consumptions were significantly higher in moderate/severe pain group than other groups ($p\text{-value} =$

0.029). On the other hand, durations of hospitalization were not different between the two groups.

Table 3 displays odds ratios of various variables associated with moderate to severe pain. The odds ratios from univariate analysis and logistic regression were interestingly different for BMI, intra-abdominal pressure, and duration of surgery. Significant odds ratio of BMI (OR 2.33, 95% CI 1.12–4.84, $p\text{-value} = 0.023$) became nonsignificant (OR 1.97, 95% CI 0.9–4.31, $p\text{-value} = 0.085$). In contrast, nonsignificant intra-abdominal pressure (OR 1.84, 95% CI 0.8–4.22, $p\text{-value} = 0.153$), and operation time (OR 1.49, 95% CI 0.86–2.61, $p\text{-value} = 0.158$) changed to be significant. Opioid use was consistently significant in univariate and multi-variate analysis.

Table 2 Analgesic use during post anesthetic care unit and postoperative period

Variables	No/mild pain (n=105)	Moderate/severe pain (n=105)	p-value
Fentanyl consumption in RR (mcg), median (IQR)	0 (0, 0)	25 (0, 50)	<0.001
Morphine consumption in RR (mg), median (IQR)	0 (0, 0)	0 (0, 2)	<0.001
Paracetamol IV in RR			
Yes	0 (0.0)	6 (100.0)	0.029
No	105 (51.5)	99 (48.5)	
NSAIDs IV in RR			
Yes	0 (0.0)	6 (100.0)	0.029
No	105 (51.5)	99 (48.5)	
Fentanyl consumption 24 hrs (mcg), median (IQR)	0 (0, 30)	0 (0, 0)	0.305
Morphine consumption 24 hrs (mg), median (IQR)	0 (0, 3)	3 (0, 12)	<0.001
Durations of hospitalization (days), median (IQR)	4 (3, 6)	4 (4, 5)	0.329

Data are presented by number (%), unless indicated otherwise

RR=recovery room, IQR=interquartile range, NSAIDs=non-steroidal anti-inflammatory drug, mcg=microgram, mg=milligram, IV=intravenous, hr=hour

Table 3 Logistic regression analyses of factors associated with moderate to severe pain after laparoscopic surgery

Factors	Crude OR (95% CI)	p-value	Adjusted OR (95% CI)	p-value
Age ≤40 years	0.88 (0.49, 1.57)	0.657	-	-
BMI ≥35 kg/m ²	2.33 (1.12, 4.84)	0.023	1.97 (0.9, 4.31)	0.085
Sex: Female	1.09 (0.61, 1.95)	0.768	-	-
ASA (Ref=class I)				
Class II	1.28 (0.42, 3.87)	0.667	-	-
Class III	1.58 (0.49, 5.12)	0.446	-	-
Type of surgery (Ref=Bariatric surgery)				
Gynecological	0.33 (0.13, 0.83)	0.018	-	-
Upper GI surgery	0.34 (0.13, 0.87)	0.025	-	-
Colorectal	0.36 (0.14, 0.94)	0.037	-	-
Urological	0.61 (0.14, 2.71)	0.519	-	-
Abdominal pressure ≥12 mmHg	1.84 (0.8, 4.22)	0.153	2.83 (1.03, 7.77)	0.04
Duration of surgery >180 mins	1.49 (0.86, 2.61)	0.158	2.23 (1.12, 4.42)	0.02
Multiple ports	1.15 (0.4, 3.31)	0.789	-	-
Local infiltration	1.42 (0.55, 3.68)	0.472	-	-
Opioid intraoperative (Ref=Fentanyl)				
Morphine	0.19 (0.04, 0.82)	0.026	0.16 (0.04, 0.71)	0.033
Paracetamol IV intraoperative	0.92 (0.42, 2.02)	0.841	-	-
NSAIDs IV intraoperative	0.81 (0.43, 1.53)	0.517	-	-

OR=odd ratio, CI=confidence interval, BMI=body mass index, ASA=American Society of Anesthesiologists, kg=kilogram, mm=millimeter, Hg=mercury, min=minute, IV=intravenous, NSAIDs=non-steroidal anti-inflammatory drugs

Discussion

In this study, we found that approximately 50.0% of the patients suffered from moderate to severe pain after laparoscopic surgery. This result was consistently same as previous literature.^{3,4} Intra-abdominal pressure equal or more than 12 mmHg and operation time more than 3 hours were significantly associated with moderate to severe pain whereas morphine as intraoperative analgesic drug significantly reduced the pain. There are several reasons for these associations.

On one hand, the intense pain in laparoscopic procedure could be determined as incisional pain and visceral pain from the effect of pneumoperitonuem.¹³ This retrospective study revealed that only abdominal pressure more than 12 mmHg and duration of surgery more than 3 hours were associated with moderate to severe pain in laparoscopic procedure. This result was similar with previous

literature.^{8,13} Akkoc et al. reported that using low insufflation pressure (10 mmHg) was associated with low postoperative pain score.¹³ Another study stated that postoperative pain score was significantly reduced in low abdominal pressure (8 mmHg) compared with standard pressure (12 mmHg) and high insufflation pressure (15 mmHg).¹⁴ According to the literature, correlation between high insufflation pressure and increased pain level is related to pneumoperitoneum effect. CO₂ pneumoperitoneum causes peritoneal stretching, overdistension of diaphragmatic muscle fiber, stretching of nerve, and releasing of inflammatory response.¹⁵ This leads to visceral pain, and shoulder pain. However, low pressure of gas would limit surgical fields with eventual difficulty in operation, though we should confirm this result with a randomized control trial study.

Operation time is another factor which impacts moderate to severe pain. Prolonged operation time (more

than 3 hours) in laparoscopic procedure increases risk of complications such as subcutaneous emphysema, and shoulder pain.¹⁶ This is the result of the prolongation of pneumoperitoneum effect.

In our study, we found that BMI \geq 35 kg/m² was not associated with pain. Several publications stated that obesity was not associated with postoperative pain.^{4,7,17} This was probably due to the pharmacokinetics of analgesia drugs.¹⁸ The metabolism of morphine was not altered in the morbidly obese when compared with normal volunteers. As a result, morphine-6-glucuronide (M6G) and morphine-3-glucuronide (M3G), the metabolites of morphine, would effectively reduce pain equally between obese and non-obese. This assumes obesity would not be associated with postoperative pain.

Morphine and fentanyl are the most commonly used analgesic drugs worldwide. Their analgesia would be useful in intraoperative and postoperative period. Similarly, in this research, morphine had more potency than fentanyl in reducing postoperative pain. Previous research has revealed that fentanyl needs more effort to control pain due to fentanyl's lipophilic nature and its tendency to redistribute after administration in adipose tissues. The short duration of fentanyl effect is due to its rapid redistribution from sites of action in the brain to sites of storage (muscle and fat).¹⁹ These pharmacokinetic and pharmacodynamic properties were not seen in morphine. Morphine is primarily metabolized by the liver uridine diphosphate glucuronosyltransferase to active M3G and M6G. These contribute to the analgesic effect and they are eliminated by renal function (Creatinine clearance).²⁰ resulting in the duration of morphine being longer than fentanyl. The limitation of this research is that it is a retrospective study with data analyzed from a single center. Additionally, there are a variety of laparoscopic surgeries that were included in this study for which the pain intensity would be different from each other.

Conclusion

Incidence of moderate to severe pain after laparoscopic surgeries is about 50.0%. Intra-abdominal pressure \geq 12 mmHg and operation time more than 3 hours are the independent risk factors of postoperative moderate and severe pain in laparoscopic surgery. Furthermore, morphine should be the appropriate opioid used in intraoperative period for reducing probability of moderate to severe pain in the postoperative period, especially in patients requiring high abdominal pressure or complicated surgery that has long duration.

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Conflict of interest

There are no potential conflicts of interests to declare.

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