

Comparison of Different Anesthetic Regimens in Patients Undergoing Laparoscopic Adjustable Gastric Banding Operations: A Prospective Randomized Trial

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Key Words: morbid obesity; laparoscopic adjustable gastric banding operation; respiratory function; remifentanyl-induced hyperalgesia; reverse Trendelenburg position.

Summary. Background and Objective. Obesity is a multisystem disorder, particularly involving the respiratory and cardiovascular systems; therefore, a multidisciplinary approach is required. In spite of widespread performance of weight reduction (bariatric) surgeries, information regarding the anesthetic care of morbidly obese patients is scarce. The aim of this study was to compare the impact of fentanyl and remifentanyl on the time of recovery, breathing parameters, and postoperative pain in morbidly obese patients undergoing laparoscopic adjustable gastric banding operations.

Material and Methods. In this prospective randomized study, 66 morbidly obese patients (BMI > 35 kg/m²), aged between 24 and 70 years, scheduled for a laparoscopic adjustable gastric banding operation were divided into 2 groups based on the opioid used for anesthesia: group 1 whose who received remifentanyl; and group 2, fentanyl). The following parameters were recorded: peripheral blood oxygenation (SpO₂) while breathing room air at baseline and 5 minutes after pre-oxygenation (100%); end-tidal carbon dioxide pressure at designated time points during the procedure; time to extubation; SpO₂ in the postanesthesia care unit; and pain intensity (using the visual analogue scale); and the presence of nausea and vomiting.

Results. The time to extubation was shorter in the remifentanyl group, but there was no significant difference in the time to discharge from the postanesthesia care unit. The recovery of respiratory parameters to the baseline values was better and faster in the remifentanyl group. The intensity of postoperative pain was similar in both groups (VAS, < 3)

Conclusions. Remifentanyl showed good analgesic properties during laparoscopic gastric banding surgery. Postanesthesia recovery and return of respiratory parameters to the baseline values was faster when remifentanyl was used. Postoperative pain and the rate of opioid-induced side effects after analgesia with remifentanyl were similar as after anesthesia with a longer acting opioid, fentanyl. Despite the problem widely discussed in literature about remifentanyl-induced hyperalgesia, no cases of analgesic overconsumption were registered in our study.

Introduction

The prevalence of morbid obesity is increasing worldwide, and surgical interventions may entail a considerable risk for obese patients. Obesity is a multisystem disorder, particularly involving the respiratory and cardiovascular systems; therefore, a multidisciplinary approach is required. In spite of widespread performance of weight reduction (bariatric) surgeries, information on the anesthetic care of morbidly obese patients is scarce.

One of many problems in the anesthetic management of morbidly obese patients (body mass index [BMI], > 35 kg/m²) is the maintenance of adequate oxygenation. Anesthesia adversely affects the respiratory function and leads to a reduced functional residual capacity (FRC). In obese patients, FRC markedly decreases with possible hypoxemia and hypercapnia during the postoperative period (1–3). The body weight is a primary factor determining arterial oxygen tension although pneumoperitoneum and position changes (Trendelenburg and reverse Trendelenburg) do not significantly alter oxygenation despite considerable deterioration in the respiratory system mechanics (3).

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Postoperative awakening and rapid recovery of spontaneous respiratory parameters depend on the agents used for anesthesia. Several studies have evaluated the benefits of the use of poorly soluble volatile anesthetic agents (desflurane and sevoflurane) in comparison with more soluble ones (isoflurane) or propofol in morbidly obese patients (4, 5). Opioids used for general anesthesia in the obese patients during laparoscopic operations vary, but ultrashort-acting opioids, such as remifentanyl and sufentanyl, are considered the opioids of choice (6). Otherwise, remifentanyl-induced hyperalgesia, especially when high doses of remifentanyl are infused, is also a matter of concern and remains a controversy (7, 8).

Because awake morbidly obese patients already have severe alterations of their respiratory mechanics (decreased chest wall and lung compliances and decreased FRC), we hypothesized that the choice of anesthetic agents would be associated with better recovery.

The aim of this prospective, randomized study was to compare the impact of fentanyl and remifentanyl on the time of recovery, breathing parameters, and postoperative pain in morbidly obese patients undergoing laparoscopic adjustable gastric banding operations.

Material and Methods

The study protocol was approved on November 6, 2008 (No. 62), by the Lithuanian Bioethics Committee and the Faculty of Medicine, Vilnius University.

After written informed consent was obtained, morbidly obese patients (BMI, >35 kg/m²) aged between 24 and 70 years, the American Society of Anesthesiologists (ASA) status class 2 and 3, and scheduled for a laparoscopic adjustable gastric banding operation (LAGB, MiniMizer extra, port medium) were included in the study. The exclusion criteria were difficult airways, obstructive pulmonary disease, ischemic heart disease, known impaired renal or hepatic function, and neuromuscular disorders.

During the preoperative visit, an anesthesiologist carried out airway assessment and laryngoscopy grading according to the Mallampati classification. The patients were classified to the following classes: class 1 the tonsils, uvula, and soft palate are clearly visible; class 2, the hard and soft palates, upper portion of tonsils, and uvula are visible; class 3, the soft and hard palates and the base of the uvula are visible; and class 4, only the hard palate is visible.

The patients were grouped using a set of computer-generated random numbers kept in sequentially numbered opaque sealed envelopes. All the patients received premedication (7.5 mg of midazolam per os at night).

On arrival to the operating room, 3-lead electrocardiography, a noninvasive measurement of blood pressure, and pulse oximetry were performed. After 5 minutes of preoxygenation with 100% oxygen, general anesthesia was induced with remifentanyl (1 μ g/kg of total body weight, TBW) during the first 45 seconds followed by an infusion of 0.1–0.15–0.2 μ g/(kg·min) of TBW (remifentanyl group) or intravenous fentanyl at a dose of 500 μ g (fentanyl group). The remifentanyl infusion rate was titrated to patients' responses. All the patients received thiopental (4 mg/kg), and tracheal intubation was facilitated with succinylcholine (1.5 mg/kg) (9). The endotracheal tube size depended on the patient's gender, i.e., female patients were intubated with size 7.5 ID, male patients with 8.5 ID. None of the patients required fiberoptic intubation. The presence of a capnography trace and bilateral breath sounds were confirmed by an anesthesiologist who performed the intubation. Anesthesia was maintained with Sevoflurane in a mixture of oxygen and air adjusted to maintain inspiratory O₂ at 70%. The mechanical ventilation was conducted with a semiclosed circuit-breathing device Dräger (tidal volume, 10 mg/kg; respiratory rate, 12–14 breaths per minute; positive end-expiratory pressure, 4 cm H₂O) to maintain the end-tidal carbon dioxide pressure (P_{ET} CO₂) between 30 and 35 mm Hg. The neuromuscular blockade was maintained with rocuronium (0.6 mg/kg ideal body weight, IBW = height in cm – 106 [for women] or 102 [for men]) (10).

After the intubation, all the patients received intramuscular diclofenac (75 mg) and intravenous dexamethasone (8 mg). Sevoflurane was maintained up to 0.8–1.2 minimum alveolar concentration, which was adjusted to keep hemodynamic changes within 20% of the baseline values measured during the preoperative anesthetic visit. The patients' lungs were ventilated with air and oxygen (FIO₂, 0.6); the initial ventilatory setting was set at the frequency of 12–14 breaths per minute, the inspiratory-to-expiratory ratio of 1:2, positive end-expiratory pressure (PEEP) of 4 cm H₂O, and tidal volume of 600–700 mL (8 mL/kg) (Dräger Ohmeda S/5 Avance). Muscle relaxation was monitored in the train-of-four values on the radial nerve; decurariation at the end of surgery was considered if the TOF values were 0.7–0.8. Entropy-based method was used to monitor the depth of anesthesia with the values kept at 50–55. Then the patient was positioned in the semisitting reverse Trendelenburg position during the entire laparoscopic procedure (pneumoperitoneum pressure was kept at 16 mm Hg) with spread legs and stretched arms. All the patients in both groups received intravenous morphine at a total dose of 10 mg 30–50 minutes before the end of the surgery.

The following parameters were recorded: peripheral blood oxygen saturation (SpO_2) while breathing room air after 5 minutes of preoxygenation with 100% oxygen via a face mask and $P_{ET}CO_2$. Muscle fasciculations due to the administration of depolarizing muscle relaxant succinylcholine were registered, and their severity was evaluated as well: no fasciculations, moderate fasciculations, and severe fasciculations. All the measurements were performed at designated time points during the procedure. The parameters of both the groups were recorded at the following time points: immediately after the intubation; 5 minutes after the intubation; at the moment when the patient's position was changed into the reverse Trendelenburg position and CO_2 insufflation was initiated; 5 minutes after the patients' position was changed into the reverse Trendelenburg position; at the time when the patient's initial position was resumed (end of the laparoscopic period of the operation); and 5 minutes after placing the patient into a neutral position.

The infusion of remifentanyl in the group 1 was discontinued when the wound was started to suture, and sevoflurane was discontinued with the last suture in both the groups. When adequate spontaneous respiration returned and verbal contact was initiated, the patients were extubated in the operating room. Then the patients were transferred to a postanesthesia care unit (PACU), they were placed in the semisitting position, and the facial mask with 5 L of oxygen per minute was applied for 1 to 1.5 hours.

In the PACU setting, SpO_2 , pain according to the visual analogue scale (VAS), and presence of nausea and vomiting were evaluated. Pain control was considered adequate if the score on the VAS was 3 and less. Rescue analgesia was maintained with intravenous morphine.

Other data recorded were the duration of anesthesia (time between induction and extubation), duration of surgery (time between incision and skin closure), additional analgesic agents in the PACU, nausea/vomiting in the PACU, time to discharge,

and SpO_2 on discharge from the PACU.

Statistical data analysis was carried out using the Statistical Package for Social Sciences (SPSS) for Windows, version 17. The intergroup differences were analyzed using the Student *t* test. To investigate the correlation between the used opioid, BMI, extubation time, and the respiratory parameters during the surgery, the Pearson correlation analysis was performed. A value of *P* less than 0.05 was considered statistically significant.

It was hypothesized that recovery would be faster in the group 1 than group 2. Therefore, the primary end-point was the time spent by the patients in the PACU. Based on the previous data from our group, it was calculated that the length of stay of 120 ± 20 minutes and a difference between the groups of 30 minutes could be considered clinically relevant. For a power of 0.9 and assuming α risk of 0.05, the sample size of 30 patients in each group was calculated to be appropriate.

Results

Altogether, 72 patients were screened during the study period from January 2009 to December 2009. The patients were assigned to either the remifentanyl group (group 1) or the fentanyl group (group 2). However, 6 patients did not meet the inclusion criteria: 4 patients were excluded because of obstructive pulmonary disease (1 patient in the group 1), and 2 patients had a history of difficult intubation (group 2). All the patients underwent the procedures uneventfully. The demographic and clinical data of the enrolled patients are presented in Table 1. There were no significant differences in the demographic data, except for age, duration of the operation or anesthesia, and SpO_2 on arrival between the groups. Hemodynamic parameters during the operation were similar in both groups (data not shown).

Table 2 shows the distribution of patients according to the Mallampati classification assessed during the preoperative visit by an anesthesiologist and severity of fasciculations. Succinylcholine was used in all the cases to facilitate tracheal intubation, and fasciculations due to the use of succinylcholine were

Table 1. Demographic and Clinical Data

Parameter	Remifentanyl Group (Group 1) n=26	Fentanyl Group (Group 2) n=40	<i>P</i>
Age, years	41.6 (10.0) [24–67]	48.3 (12.9) [26–70]	0.02
Sex, male/female, n	8/18	14/26	0.59
BMI, kg/m ²	45.9 (6.9) [35–61]	48.3 (7.0) [35–67]	0.19
Duration of surgery, min	69.8 (16.5) [45–130]	69.3 (18.8) [40–125]	0.9
Duration of anesthesia, min	90 (19.5) [50–150]	99.9 (22.1) [65–155]	0.06
SpO_2 on arrival, %	95.8 (2.3) [90–100]	95.1 (2.3) [88–99]	0.25

Values are mean (SD) [range] unless otherwise noted.

Table 2. Distribution of Patients According to the Mallampati Classification and Severity of Fasciculations

Variable	Remifentanil Group (Group 1) n=26	Fentanyl Group (Group 2) n=40
Mallampati class		
1	2 (7.7)	2 (5.0)
2	10 (38.5)	9 (22.5)
3	10 (38.5)	13 (32.5)
4	4 (15.4)	16 (40)
Fasciculation		
Absent	13 (50.0)	18 (22.2)
Moderate	11 (42.3)	20 (20.0)
Severe	2 (7.7)	2 (5.0)

Values are number (percentage).

registered. There were only a few cases of failed intubation in both the groups: 2 patients in the group 1 were intubated from the third attempt, 1 patient and 2 patients in the group 2 were intubated from the second and third attempts, respectively. There was no significant difference comparing the severity of fasciculations between the groups ($P=0.8$).

Respiratory parameters were monitored during the operation and were registered at 6 different time points. Fig. 1 represents the change in $P_{ET}CO_2$ during the operation at the particular time points. No significant differences were registered comparing the groups. Among the morbidly obese patients, there was a correlation between BMI and $P_{ET}CO_2$ when the patients were placed in the reverse Trendelenburg position ($r=0.264$, $P=0.018$) (Fig. 2) and after 5 minutes ($r=0.378$, $P=0.03$).

According to the protocol, all the patients spent 120 minutes in the PACU and were discharged. Before the discharge, the patients were breathing room air for at least 20 minutes. SpO_2 was registered on discharge, and any complaints of breathing shortage were taken into account. There were no patients in both the groups who needed extra time to spend in the PACU (Table 3).

Peripheral blood oxygenation while breathing room air was registered on patients' arrival to the operating room and on discharge. There was no significant correlation between BMI and SpO_2 on arrival ($r=-0.337$, $P=0.25$), but on discharge, BMI and SpO_2 were significantly correlated ($r=-0.269$, $P=0.016$) (Fig. 3).

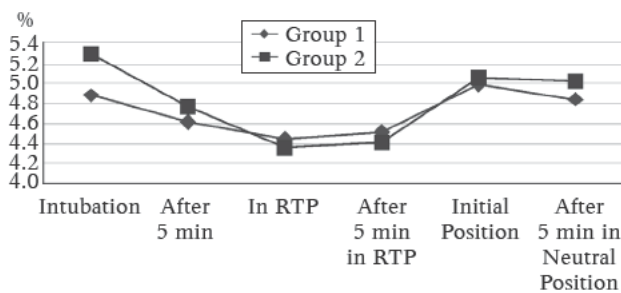


Fig. 1. Intraoperative changes in the end-tidal carbon dioxide pressure ($P_{ET}CO_2$)

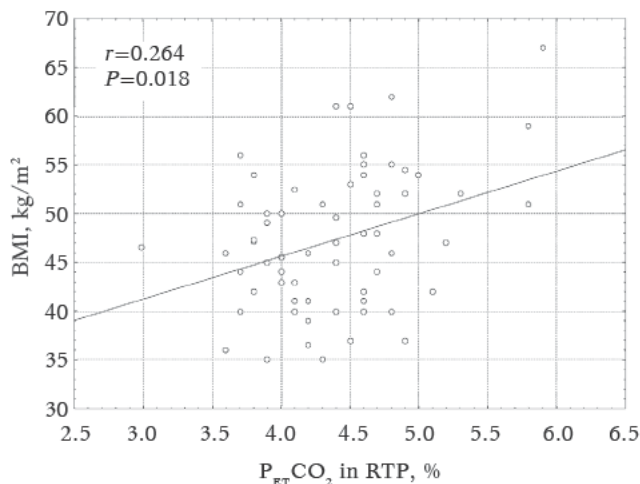


Fig. 2. Correlation between body mass index (BMI) and end-tidal carbon dioxide pressure ($P_{ET}CO_2$) in the reverse Trendelenburg position (RTP)

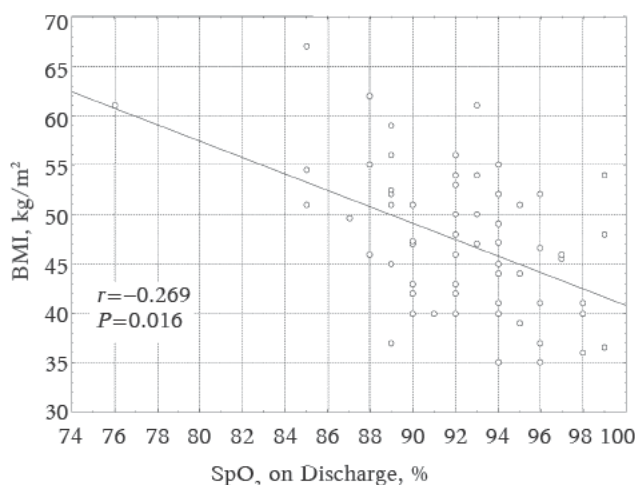


Fig. 3. Correlation between body mass index (BMI) and peripheral blood oxygen saturation (SpO_2) on discharge

Table 3. Postoperative Data: Time to Extubation, Additional Analgesic Consumption, and Nausea/Vomiting in the Postanesthesia Care Unit

Parameter	Remifentanil Group (Group 1) n=26	Fentanyl Group (Group 2) n=40	P
Time to extubation, min	6.5 (3.4) [5–20]	14.3 (7.6) [5–30]	0.001
Additional analgesic (morphine), mg	12.1 (4.7) [10–25]	10.5 (2.5) [5–20]	0.12
Nausea/vomiting in the PACU, n (%)	2 (7.6)	4 (10)	0.19
SpO_2 on discharge, %	94.1 (3.1) [88–99]	90.9 (4.0) [82–99]	0.00045

Values are mean (SD) (range) unless otherwise noted. PACU, Postanesthesia Care Unit; SpO_2 , peripheral blood oxygen saturation.

On arrival, there was no difference in SpO₂ between the groups ($P=0.7$), but on discharge, the group 1 showed significantly better SpO₂ values than the group 2 ($P<0.00045$). The patients receiving remifentanyl (group 1) showed significantly faster recovery after the operation and a shorter time to extubation, but the opioid (remifentanyl or fentanyl) used for analgesia during the operation had no impact on the patients' stay time in the PACU.

Discussion

The results of this study indicated that the time to extubation was shorter in the remifentanyl group, but the time to discharge from the PACU after laparoscopic adjustable gastric banding operations was not influenced by the choice of the opioid administered during the operation. The length of stay in the PACU was similar in both groups, and the return of respiratory parameters to the preoperative values was better and faster in the remifentanyl group. Postoperative pain according to the VAS was similar in both the groups.

Succinylcholine was used to facilitate tracheal intubation for its rapid onset of action and short duration of action because in morbidly obese patients, hemoglobin desaturation occurs rapidly after apnea, and the intubation of trachea must be accomplished quickly (9). The evaluation of morbidly obese patients according to the Mallampati classification loses its value in predicting the difficulty of forthcoming tracheal intubation; 23 and 20 patients were classified as Mallampati class 3 and 4, respectively, but only 2 patients classified as Mallampati class 3 and 4 were intubated after repeated attempts, the same as 2 patients classified as Mallampati class 1 and 2.

The present study considered the postoperative recovery time, time to extubation, postoperative pain, and episodes of postoperative nausea and vomiting. Postoperative pain according to the VAS in the PACU was similar. Morphine as a rescue analgesic was used in both the groups, and although greater usage was documented in the remifentanyl group, the difference between the groups was insignificant. Besides, a laparoscopic gastric banding operation is not considered a particularly painful procedure. In addition, nonsteroidal anti-inflammatory therapy (Diclac, 75 mg intravenously) was initiated during the anesthesia.

In morbidly obese patients, alfentanil, fentanyl, and remifentanyl can be safely used, but a study by Gaszynski et al. reported a higher frequency of postoperative nausea and vomiting and more severe postoperative pain in the remifentanyl group (11). The reason for a larger discrepancy in the postoperative need of a rescue analgesic between remifentanyl and other opioids is unknown. However, there are several possible explanations: pa-

tients receiving remifentanyl develop acute opioid tolerance while other opioids have longer acting postoperative residual effects, or remifentanyl causes hyperalgesia (12). Moreover, the evidence of dose-dependent hyperalgesia exists, and a number of articles describe additional options how to solve this problem (13, 14). Additives, such as magnesium sulfate, ketamine, or dexmetomedine, can decrease the pain threshold at the peri-incisional areas (13, 14), but ketamine may cause other well-known problems like hallucinations and an altered body image in the early postoperative period. Besides, the infusion dose of remifentanyl also matters: the higher the individual requirements are, the higher the risk of postoperative hyperalgesia appears because acute opioid tolerance seems to be dose dependant (15).

Respiratory mechanics in healthy, normal-weight, nonanesthetized patients changes minimally over a wide range of postures, including both 30° head-up and head-down positions. In healthy non-obese patients, abdominal insufflation in the head-down position only moderately decreases the respiratory system compliance. Morbid obesity itself may cause a significant respiratory system alteration (16–18). The respiratory system compliance exponentially decreases as a function of the increased body mass index (2). A reduced lung volume can explain the alterations of respiratory system mechanics (decreased compliance and increased resistance) in anesthetized morbidly obese patients in the supine position. Induction of 16-mm Hg CO₂ pneumoperitoneum and positioning of these patients in the reverse Trendelenburg posture (beach chair position) reduced the respiratory compliance and increased the resistance (19). Similar studies revealed a correlation between body mass index and inspiratory resistance in the presence of pneumoperitoneum while performing laparoscopic operations in the reverse Trendelenburg position (2). The hypothesis that the reverse Trendelenburg position will improve respiratory mechanics because of gravitational effects on abdominal contents has been proven, but only in case of open bariatric surgery. In case of laparoscopic adjustable gastric banding operations with CO₂ insufflation, the reverse Trendelenburg position had no beneficial effect on respiratory parameters, and our findings are in agreement with other studies (20).

Some limitations of our study should be mentioned. The study had to include 80 patients had a strict time to be completed. Unfortunately, in 2009, there were problems with the delivery of remifentanyl, and this led to the different sample sizes in the groups. Despite this, our study succeeded to highlight some substantial differences between the patients receiving remifentanyl and those receiving fentanyl.

Conclusions

Remifentanil showed good analgesic properties during laparoscopic gastric banding surgery. Postanesthesia recovery and return of respiratory parameters to the baseline values was faster when remifentanil was used. Postoperative pain and the rate of opioid-induced side effects after analgesia with remifentanil were similar as after anesthe-

sia with a longer acting opioid, fentanyl. Despite the problem widely discussed in literature about remifentanil-induced hyperalgesia, no cases of analgesic overconsumption were registered in our study.

Statement of Conflict of Interest

The authors state no conflict of interest.

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Received 3 November 2011, accepted 30 October 2012