

Comparison of respiratory mechanics between sevoflurane and propofol-remifentanil anesthesia for laparoscopic colectomy

Si Ra Bang¹, Sang Eun Lee¹, Hyun Joo Ahn², Jie Ae Kim², Byung Seop Shin², Hee Jin Roe², and Woo Seog Sim²

Department of Anesthesiology and Pain Medicine, ¹Haeundae Paik Hospital, Inje University School of Medicine, Busan, ²Samsung Medical Center, Sungkyunkwan University School of Medicine, Seoul, Korea

Background: The creation of pneumoperitoneum and Trendelenburg positioning during laparoscopic surgery are associated with respiratory changes. We aimed to compare respiratory mechanics while using intravenous propofol and remifentanil vs. sevoflurane during laparoscopic colectomy.

Methods: Sixty patients undergoing laparoscopic colectomy were randomly allocated to one of the two groups: group PR (propofol-remifentanil group; n = 30), and group S (sevoflurane group; n = 30). Peak inspiratory pressure (PIP), dynamic lung compliance (C_{dyn}), and respiratory resistance (R_{rs}) values at five different time points: 5 minutes after induction of anesthesia (supine position, T1), 3 minutes after pneumoperitoneum (lithotomy position, T2), 3 minutes after pneumoperitoneum while in the lithotomy-Trendelenburg position (T3), 30 minutes after pneumoperitoneum (T4), and 3 minutes after deflation of pneumoperitoneum (T5).

Results: In both groups, there were significant increases in PIP and R_{rs} while C_{dyn} decreased at times T2, T3, and T4 compared to T1 (P < 0.001). The R_{rs} of group PR for T2, T3, and T4 were significantly higher than those measured in group S for the corresponding time points (P < 0.05).

Conclusions: Respiratory mechanics can be adversely affected during laparoscopic colectomy. Respiratory resistance was significantly higher during propofol-remifentanil anesthesia than sevoflurane anesthesia. (Korean J Anesthesiol 2014; 66: 131-135)

Key Words: Laparoscopy, Propofol, Remifentanil, Respiratory mechanics, Sevoflurane.

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Corresponding author: Woo Seog Sim, M.D., Ph.D., Department of Anesthesiology and Pain Medicine, Samsung Medical Center, Sungkyunkwan University School of Medicine, 50, Ilwon-dong, Gangnam-gu, Seoul 135-710, Korea. Tel: 82-2-3410-0356, Fax: 82-2-3410-0361, E-mail: anesthe@skku.edu

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Introduction

Laparoscopic colectomy is increasingly being performed because it is minimally invasive technique and therefore associated with decreased blood loss, a shorter hospital stay and fewer wound problems than open colectomy [1,2]. However, the creation of pneumoperitoneum and Trendelenburg positioning for this procedure are associated with an increase in airway pressure and a decrease in lung compliance, as well as barotrauma [3]. These alterations in respiratory mechanics could adversely affect patient outcomes.

Both total intravenous anesthesia with propofol-remifentanyl and inhalation anesthesia with sevoflurane are widely used. Generally, both sevoflurane and propofol can decrease respiratory resistance (Rrs) or peak inspiratory pressure (PIP), and/or increase dynamic compliance (Cdyn) [4-13]. In contrast, opioids can induce chest wall rigidity but effectively be reversed by neuromuscular blocking agents [14,15]. A previous experimental study concluded that propofol assured better respiratory conditions than sevoflurane during laparoscopic surgery [4]. However, that study was conducted using a porcine model.

Our purpose in this study was to compare the respiratory mechanics of patients anesthetized by intravenous propofol-remifentanyl vs. inhalant sevoflurane during laparoscopic colectomy.

Materials and Methods

This study was performed after approval from the local institutional review board. Written informed consent was obtained from all patients. Sixty patients, aged between 40 and 75 years old, with American Society of Anesthesiologists physical status I and II who were scheduled to undergo laparoscopic colectomy were enrolled from August 2009 to December 2009. Patients with cardiovascular disease, pulmonary disease, morbid obesity (>20% heavier than ideal body weight), or who were current smokers were excluded from the study.

Patients were allocated randomly to one of two groups: group PR (propofol-remifentanyl group; n = 30), or group S (Sevoflurane group; n = 30) using a computer-generated table of random numbers. Patients were not premedicated. Hemodynamics were monitored standardly with non-invasive blood pressure, oxygen saturation and electrocardiography.

To induce anesthesia, all patients were ventilated with 100% O₂ in a mask. In group PR, after intravenous injection of 20 mg lidocaine, propofol (Fresofol MCT 2% Inj., Fresenius Kabi, Graz, Austria) and remifentanyl hydrochloride (Ultiva[®], Glaxo-SmithKline, Parma, Italy) were administered at 5 µg/ml and 3 ng/ml, respectively, using an effect-site concentration target-controlled infusion pump (Orchestra[®], Fresenius Vial, Brezins,

France). In group S, anesthesia was induced with propofol (Pofol inj., Jeil Pharm CO., Seoul, Korea) 2 mg/kg, fentanyl 1 µg/kg and sevoflurane. All patients were intubated with a 7.5 mm internal diameter endotracheal tube at about 3 min after injection of 0.8 mg/kg rocuronium.

During maintenance of anesthesia, propofol and remifentanyl infusions or sevoflurane were titrated to maintain a mean blood pressure within 20% of baseline. All patients were ventilated with a fixed protocol: a tidal volume of about 8 ml/kg, 12 breaths/minute, and an inspiratory-to-expiratory ratio of 1 : 2 in 1.5 L/min of O₂-air mixture at FiO₂ 0.5 using a ventilator (Aestiva/5, Datex Ohmeda, Helsinki, Finland). Muscle relaxation was maintained between 90–95% in response to a train of four stimulations with continuous infusion of 0.05 mg/kg/hr vecuronium. Patients who were injected with any other drugs for hemodynamic control or anesthesia, or whose position changed for any reason before data were collected, were excluded from this study.

A disposable spirometry tube was connected to the proximal end of the tracheal tube. Respiratory mechanics were displayed and measured using an anesthesia monitor (Datex Ohmeda S/5TM compact anesthesia monitor, General electric, Helsinki, Finland).

Intraabdominal CO₂ pressure was maintained at around 12 mmHg during pneumoperitoneum in a lithotomy position. The operative table had a head-down tilt of 20 degrees to place the patient in the Trendelenburg position. Respiratory mechanical parameters (PIP, Cdyn and Rrs) were recorded at five different time points: 5 minutes after induction of anesthesia (supine position, T1), 3 minutes after pneumoperitoneum (lithotomy position, T2), 3 minutes after pneumoperitoneum while in the lithotomy-Trendelenburg position (T3), 30 minutes after pneumoperitoneum (T4), and 3 minutes after deflation of pneumoperitoneum (T5). We calculated the mean value of data measured three times during three consecutive ventilations during periods when the compliance loops were not distorted by any interference, such as instrument changes.

Statistical analysis was performed with the software program SAS 9.1.3 (SAS Institute, Inc., Cary, NC, USA). Sample size was determined based on a preliminary study. To be able to measure a 15% difference in PIP between two groups with a power of 80% and an alpha error of 0.05, we calculated that the minimum sample size required was 28 patients per group. We therefore included 30 patients in each group to account for dropout.

Demographic data were compared using a two-sample t-test or chi-square test. Changes in respiratory mechanic parameter values with time were analyzed by repeated measures analysis of variance (ANOVA) with Bonferroni correction for post hoc comparisons. Data are reported as mean ± SD. A P value < 0.05 was considered statistically significant.

Results

Patient characteristics were not different between the two groups (Table 1). All patients were hemodynamically stable. No episodes of desaturation occurred.

In all groups, there was a significant increase in PIP and Rrs, and a decrease in Cdyn at T2, T3, and T4 compared to T1 ($P < 0.001$) (Table 2). Respiratory mechanic parameters were not significantly different at T3 and T4 compared to T2, while those at T5 were restored to values similar to those measured at T1 (Table 2).

PIP and Cdyn were not significantly different between the two groups at any time point (Table 2). Only Rrs was significantly higher during propofol-remifentanyl anesthesia than during sevoflurane anesthesia at T2, T3, and T4 ($P < 0.05$) (Table 2).

Table 1. Patient Characteristics, Duration of Anesthesia and Surgery

	Group PR (n = 30)	Group S (n = 30)
Sex (M/F)	16/14	18/12
Age (yr)	59.4 ± 10.9	54.3 ± 10.6
Height (cm)	160.8 ± 10.3	160.2 ± 7.8
Weight (kg)	63.5 ± 9.8	61.5 ± 8.6
Body mass index (kg/m ²)	24.5 ± 2.4	23.9 ± 2.5
ASA I/II (n)	13/17	16/14
Anesthetic time (min)	189.7 ± 53.3	204.0 ± 58.5
Operation time (min)	152.1 ± 53.3	165.5 ± 59.2

Data are expressed as number or mean ± standard deviation. Group PR: propofol-remifentanyl group, Group S: sevoflurane group.

Table 2. Changes in Respiratory Mechanic Parameters at Different Times

Variable	Situation	Group PR	Group S
Peak inspiratory pressure (cmH ₂ O)	T1	15.5 ± 3.0	14.6 ± 2.8
	T2	24.4 ± 5.1*	23.2 ± 5.0*
	T3	25.5 ± 4.8*	23.8 ± 5.2*
	T4	27.1 ± 5.1*	25.4 ± 4.6*
	T5	17.8 ± 3.4	17.8 ± 3.3
Dynamic compliance (ml/cmH ₂ O)	T1	63.3 ± 15.6	63.2 ± 14.5
	T2	33.6 ± 10.1*	35.5 ± 12.7*
	T3	31.3 ± 9.3*	33.4 ± 11.4*
	T4	30.2 ± 10.3*	30.4 ± 7.2*
	T5	51.4 ± 12.1	51.0 ± 13.4
Respiratory resistance (ml/cmH ₂ O/s)	T1	9.8 ± 2.9	8.4 ± 2.9
	T2	12.7 ± 5.6*. [†]	10.6 ± 4.0*. [†]
	T3	12.9 ± 5.3*. [†]	10.5 ± 3.2*. [†]
	T4	12.0 ± 4.5*. [†]	10.3 ± 3.4*. [†]
	T5	8.5 ± 2.4	7.5 ± 1.7

All values are expressed as mean ± standard deviation. Group PR: propofol-remifentanyl group, Group S: sevoflurane group. T1: 5 minutes after induction of anesthesia (supine position), T2: 3 minutes after pneumoperitoneum (lithotomy position), T3: 3 minutes after pneumoperitoneum while in the lithotomy-Trendelenburg position, T4: 30 minutes after pneumoperitoneum, T5: 3 minutes after deflation of pneumoperitoneum. * $P < 0.001$ compared to T1 value in each group. [†] $P < 0.05$ compared between groups.

Discussion

The use of intravenous propofol-remifentanyl or inhalant sevoflurane during laparoscopic colectomy was associated with an increase in PIP and Rrs a decrease in Cdyn values during the creation of pneumoperitoneum and when patients were in the Trendelenburg position. Patients in group PR had significantly higher respiratory resistance values than those in group S during the creation of pneumoperitoneum and placement in the Trendelenburg position.

We measured respiratory mechanic parameters using a computed side stream monitor. Lung compliance (C) and airway resistance (Raw) were calculated for each breath from the following equation: $C = TV_{exp}/(P_{plat}-PEEP)$ and $P(t) = Raw \, dV/dt + V(t)/C + PEEP$, where TV_{exp} = expiratory tidal volume, P_{plat} = end-inspiratory lung pressure, $PEEP$ = positive end-expiratory lung pressure, and $P(t)$ and $V(t)$ = pressure and volume at a certain time (t), respectively. Cdyn and Rrs measured in this study were not the exact C and Raw, but total compliance and resistance values that included resistance and elastic forces from the chest wall and the lung tissue [6,16-18]. However, the changes in total resistance and compliance reflect Raw and C [6,19] and the effects of the chest wall and lung tissue would be little or unchanged, equally in all patients; therefore, this is unlikely to change our results. The advantage of using this technique is that data are displayed continuously; furthermore, these values have been shown to be accurate in previous studies [5-7,11,16]. Oikkonen and Tallgren [16] reported that Cdyn decreased immediately after insufflation and was not influenced by duration of pressurization, so the timing of our measurements of respiratory parameters was appropriate. Furthermore, data were mean values of three measurements taken when the displayed loop was not distorted.

During laparoscopy, peritoneal CO₂ insufflation increases intra-abdominal pressure, elevates the diaphragm, and increases intra-thoracic pressure [20]. This leads to an increase in PIP and Raw as well as a decrease in functional residual capacity and pulmonary compliance [5,16,20-23]. Oikkonen and Tallgren [16] reported that there was no association between age, sex, BMI, duration of pressurization, or changes in Cdyn during laparoscopic surgery. Some studies have reported that the degree of muscle relaxation and smoking history have no significant effect on respiratory mechanics during mechanical ventilation [6,17,24]. However, other studies have reported that several factors (patient's position, BMI, anesthetic agents etc.) affect the extent of respiratory mechanics changes during laparoscopic surgery [5,16,20,22,23]. These changes could be aggravated in patients with pre-existing pulmonary diseases, or morbid obesity [21,22], which is why we excluded these patients. In our study, there were no significant differences between the two groups

with respect to these factors; the only difference was the type of anesthetic agent used.

Sevoflurane is known to act as a bronchodilator by directly relaxing bronchial smooth muscle and inhibiting vagal airway reflexes, similar to other inhalation anesthetics [4-9]. The mechanism of the direct relaxing effect of inhalation anesthetics on airway smooth muscle may involve a reduction of Ca^{2+} concentration and perturbation of calcium homeostasis [9,19]. Sevoflurane may also have a protective effect against airway constriction [7,9]. Sivaci et al. [5] reported that sevoflurane significantly reduced Rrs in laparoscopic surgery.

Propofol has also been reported to prevent bronchoconstriction and to induce bronchodilatation under mechanical ventilation by anticholinergic effects on the airway [11-13]. Propofol also has direct relaxant effects on airway smooth muscle [10,13,25]. Although the exact mechanism of action of propofol is still unknown, it has been postulated that it reduces the influx of calcium ions in vitro [13,25].

Opioids increase pulmonary resistance and decrease pulmonary compliance. These changes are not caused by a change in the autonomic nervous system or airway smooth muscle, but spasm of the chest wall muscles that can be abolished by neuromuscular blockers [14,15]. Intravenous lidocaine has been shown to have bronchodilatory effects in vitro and in vivo [17,26]. However, it has no bronchodilatory effect at clinical concentrations, and does not protect against bronchoconstriction [17,27]. Thus infusion of remifentanyl and 20 mg lidocaine in group PR may have had little influence on respiratory mechanics in our study.

To the best of our knowledge, only one previous report has compared respiratory mechanics under propofol and sevoflurane anesthesia during laparoscopy; the present study represents the first clinical study. Puglisi et al. [4] reported that sevoflurane resulted in significantly higher airway pressure than propofol during laparoscopy in a porcine model. They argued that propofol had a stronger relaxant action on the airway muscles than sevoflurane. We did not find any significant differences in airway pressure between the two groups. There are several reasons for this discrepancy. First, we used remifentanyl in group PR only, whereas Puglisi et al. used fentanyl in both groups. We cannot exclude the possibility that remifentanyl may have affected pulmonary mechanics in group PR, because we did not compare chest wall rigidity or extent of chest wall relaxation between the two groups. However, the influence of remifentanyl on respiratory mechanics is likely to have been negligible, because we continuously infused muscle relaxant.

Second, there were differences in the amount of anesthetics used between our study and the previous study. Puglisi et al. used 12 mg/kg/hr propofol, while we used approximately 8 mg/kg/hr in our study. We were not able to directly compare the amount of anesthetics because they did not specify the concentration of sevoflurane they used. However, because we did not use remifentanyl in group S, we might have used relatively higher concentration of sevoflurane in proportion to propofol than Puglisi and colleagues. Although there is controversy about whether inhalation anesthetics have a dose-dependent bronchodilatory effect [7,8], relatively higher concentration of sevoflurane could explain the differences in results between the two studies. The exact mechanisms of airway relaxation by propofol and sevoflurane are still unknown. However, sevoflurane acts more by direct airway smooth muscle relaxation than vagolysis, while propofol acts the other way [4,17,25,28]. Although the exact mechanisms and bronchodilatory effects of propofol and sevoflurane have yet to be determined, sevoflurane might be a better bronchodilator than propofol.

Third, propofol may have reduced PIP by relaxing the chest wall muscle to a greater extent than sevoflurane in Puglisi and colleagues' study. Both sevoflurane and propofol are known to relax skeletal muscles [29,30]. If propofol is a stronger skeletal muscle relaxant than sevoflurane or used at a relatively higher dose than sevoflurane, it could have relaxed the chest wall muscle more effectively. If Puglisi et al. did not use muscle relaxant, this effect could have been exaggerated, resulting in a decrease in PIP greater than that observed for sevoflurane, in contrast to our study findings.

The limitations of the current study are listed below. This study was a single-blinded study. Furthermore, lack of invasive monitoring makes precise interpretation of respiratory and cardiovascular effects impossible. We did not compare propofol and sevoflurane directly; rather, we compared total intravenous anesthesia and sevoflurane, as this more closely reflects the clinical situation.

Respiratory mechanics can be adversely affected during laparoscopic colectomy. Among them, Rrs increased significantly more during propofol-remifentanyl anesthesia than during sevoflurane anesthesia. Further controlled studies in patients with limited pulmonary reserves due to obesity or chronic obstructive pulmonary disease are needed.

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