

Intraoperative Recruitment Maneuver Reverses Detrimental Pneumoperitoneum-induced Respiratory Effects in Healthy Weight and Obese Patients Undergoing Laparoscopy

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ABSTRACT

Background: Pulmonary function is impaired during pneumoperitoneum mainly as a result of atelectasis formation. We studied the effects of 10 cm H₂O of positive end-expiratory pressure (PEEP) and PEEP followed by a recruitment maneuver (PEEP+RM) on end-expiratory lung volume (EELV), oxygenation and respiratory mechanics in patients undergoing laparoscopic surgery.

Methods: Sixty consecutive adult patients (30 obese, 30 healthy weight) in reverse Trendelenburg position were prospectively studied. EELV, static elastance of the respiratory system, dead space, and gas exchange were measured before and after pneumoperitoneum insufflation with zero end-expiratory pressure, with PEEP alone, and with PEEP+RM. Results are presented as mean ± SD.

Results: Pneumoperitoneum reduced EELV (healthy weight, 1195 ± 405 vs. 1724 ± 774 ml; obese, 751 ± 258 vs. 886 ± 284 ml) and worsened static elastance and dead space in both groups (in all $P < 0.01$ vs. zero-end expiratory pressure before pneumoperitoneum) whereas oxygenation was

unaffected. PEEP increased EELV (healthy weight, 570 ml, $P < 0.01$; obese, 364 ml, $P < 0.01$) with no effect on oxygenation. Compared with PEEP alone, EELV and static elastance were further improved after RM in both groups ($P < 0.05$), as was oxygenation ($P < 0.01$). In all patients, RM-induced change in EELV was 16% ($P = 0.04$). These improvements were maintained 30 min after RM. RM-induced changes in EELV correlated with change in oxygenation ($r = 0.42$, $P < 0.01$).

Conclusion: RM combined with 10 cm H₂O of PEEP improved EELV, respiratory mechanics, and oxygenation during pneumoperitoneum whereas PEEP alone did not.

What We Already Know about This Topic

- ❖ Pulmonary function is impaired during laparoscopy as a result of diaphragm shift, which causes airway closure and collapse of dependent lung regions.

What This Article Tells Us That Is New

- ❖ In patients undergoing laparoscopy, positive end-expiratory pressure increased respiratory elastance but did not improve oxygenation. Addition of a recruitment maneuver increased respiratory elastance and oxygenation in normal weight and obese patients.

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GENERAL anesthesia, even in patients with healthy lungs, causes impairment in gas-exchange and respiratory mechanics.¹ Such effects are mainly the result of the formation of atelectasis.^{2,3} Laparoscopic surgery worsens respiratory mechanics in healthy weight⁴⁻⁶ and obese patients.⁷⁻¹¹ The increase in intraabdominal pressure during pneumoperitoneum causes cranial shift of the end-expiratory position of the diaphragm, further reducing end-expiratory lung volume (EELV), and predisposes patients to airway closure and collapse of dependent lung regions.¹²⁻¹⁵

Several strategies have been investigated to improve oxygenation and respiratory mechanics during laparoscopic surgery. The use of high positive end-expiratory pressure

(PEEP) and positioning (e.g., beach chair position) has effectively improved respiratory function and oxygenation in obese patients^{7–11} whereas PEEP alone improved oxygenation only slightly.^{16,17} PEEP also efficiently counteracted the upward shift of the diaphragm during laparoscopy in healthy weight patients, limited surgical effects on respiratory mechanics, and improved oxygenation.^{18–20}

The use of a recruitment maneuver (RM) effectively re-expanded atelectasis after anesthesia induction, increased EELV, and improved oxygenation in healthy weight^{21,22} and obese patients.²³ RMs also improved oxygenation in obese patients undergoing laparoscopic procedures.^{17,24,25} However, a single RM may not be sufficient,^{23,24} and PEEP is required to prevent rapid reoccurrence of atelectasis—especially when a high-inspired oxygen fraction is used.²⁶ Conversely, although pneumoperitoneum led to increased atelectasis and altered respiratory mechanics in healthy weight patients,^{5,27} the value of RMs in such patients has been little studied during laparoscopy.²⁸

Most of the aforementioned studies investigated the effects of PEEP and RM on oxygenation. However, alveolar recruitment—an anatomical phenomenon that exhibits as restored aeration on computed tomography—often fails to coincide with functional recruitment as defined by improved gas exchange.²⁹ In addition, Strang *et al.*²⁷ recently showed that oxygenation did not correlate with atelectasis formation during pneumoperitoneum, indicating that oxygenation may be a poor indicator of the extent of lung collapse.^{12,27} Oxygenation may, therefore, be an unreliable marker of recruitment effects²⁸ and inadequately sensitive to detect lung overdistension.^{30,31} Despite the potential clinical relevance thereof, intraoperative measurement of functional residual capacity is uncommon in routine practice.^{32,33} A functional residual capacity value—defined as the relaxed equilibrium volume of the lungs when there is no pressure difference between the alveoli and the atmosphere—is normally obtained in a spontaneously breathing patient at the end of a normal expiration. In this context, EELV is used to denote functional residual capacity during mechanical ventilation. We previously demonstrated that, after induction of anesthesia, PEEP improves efficiently both EELV and respiratory mechanics, with no major effect on oxygenation.³⁴ Ventilation at low EELV may instigate or worsen lung injury, possibly as a result of repeated airway closure.³⁵ In addition, EELV was found to be a more sensitive indicator of PEEP-induced re-aeration and alveolar recruitment than was oxygenation.³⁶ Therefore, we investigated the effects of RM after application of PEEP, on EELV modifications, respiratory mechanics, and oxygenation in healthy weight and obese patients undergoing laparoscopic surgery. We hypothesized that RM would be useful to counteract the detrimental effects of pneumoperitoneum, especially after EELV reduction.

Materials and Methods

After obtaining the approval of our institutional review board (Clermont-Ferrand, France), written informed con-

sent was obtained from all patients. Sixty adult patients (30 obese [body mass index higher than 35 kg/m²], 30 healthy weight [body mass index less than 25 kg/m²]) with American Society of Anesthesiologists Physical Status Classification scores of 1–3, scheduled for laparoscopic procedures of at least 1 h, were prospectively included in the current study. Exclusion criteria were age younger than 18 yr, pregnancy, emergency surgery, heart failure (defined as New York Heart Association classification more than 3), coronary disease, and chronic obstructive pulmonary disease.

Anesthetic management was standardized as follows. General anesthesia was induced using propofol (2 mg/kg) and remifentanyl (0.25 $\mu\text{g} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$) and was maintained, to a target bispectral index between 40 and 50, with continuous infusion of propofol (using a target-controlled infusion) and remifentanyl. Anesthetic concentrations were based on ideal body weight. Muscle paralysis was induced with succinylcholine (1 mg/kg) to facilitate tracheal intubation (cuffed tube 7–7.5; Portex, Inc., London, England), and was maintained with subsequent bolus doses of cisatracurium as indicated by orbicular nerve stimulation (train-of-four ratio). No patient required fiber-optic intubation. The duration of anesthesia induction (defined as the time between the end of preoxygenation and tracheal intubation) and occurrence of hypoxic apnea (defined as peripheral oxygen saturation levels less than 92%³⁷) after anesthesia was induced were recorded for all patients. Intraoperative fluid intake was maintained using 8 ml $\cdot \text{kg}^{-1} \cdot \text{h}^{-1}$ of normal saline solution. Standard monitoring included continuous electrocardiograph, heart rate, peripheral oxygen saturation, and end-tidal carbon dioxide concentration (ETCO₂) recording. The radial artery was cannulated before induction of anesthesia, in line with the standard practice of our institution, for invasive blood pressure and blood gas monitoring. Arterial pulse pressure variation (ΔPP) was monitored throughout the surgical procedure, as previously described.³⁸ Bolus doses of hydroxyethylstarch (HES 130/0.4, Voluven; Fresenius Kabi, Bad Hamburg, Germany) were given, as necessary, up to 50 ml/kg, to maintain ΔPP at less than 13%, especially before RM.

Study Protocol

A schema of the protocol is shown in figure 1. In both weight groups, anesthesia induction and the study procedure were performed in the beach chair position, as previously described.⁷ Before induction of anesthesia, preoxygenation (spontaneous breathing of 100% oxygen *via* facemask) was conducted for 5 min before tracheal intubation. Immediately after intubation, patients were mechanically ventilated (Engström Carestation; Datex-Ohmeda, General Electric, Helsinki, Finland) with the ventilator in the volume-controlled mode and tidal volume at 8 ml/kg⁻¹ ideal body weight, a respiratory rate adjusted to maintain an arterial carbon dioxide tension of 35–42 mmHg, and an inspiratory/expiratory ratio of 1/2. The inspiratory oxygen fraction (FiO₂) was 0.5

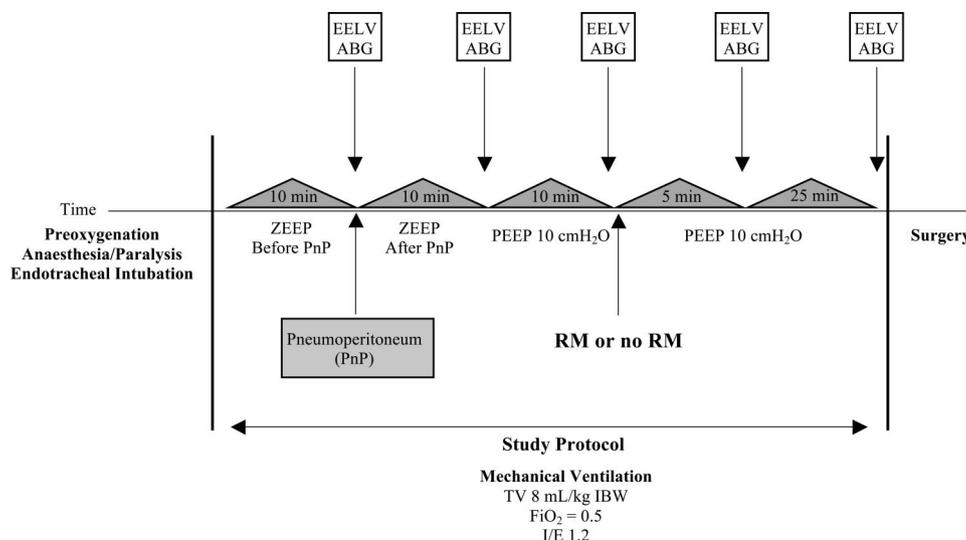


Fig. 1. Schema of the procedure. Measurements were obtained: (1) before pneumoperitoneum (PnP) insufflation with zero end-expiratory pressure (ZEEP), (2) 10 min after PnP insufflation with ZEEP, (3) 10 min after positive end-expiratory pressure (PEEP) 10 cm H₂O was applied, (4) 5 min, and then (5) 30 min after intervention ([recruitment maneuver] or no RM). Apart from application of recruitment maneuver and adjustment of PEEP level, basal ventilatory settings were kept constant throughout. ABG = arterial blood gas; EELV = end-expiratory lung volume; FiO₂ = inspiratory oxygen fraction; IBW = ideal body weight; I/E = inspiratory-expiratory ratio; TV = tidal volume.

and zero end-expiratory pressure (ZEEP) was applied before the onset of pneumoperitoneum (step 1).

Pneumoperitoneum was generated by insufflating carbon dioxide into the abdomen with the intraabdominal pressure maintained at 12 cm H₂O (40-l High-Flow Insufflator; Stryker Endoscopy, San Jose, CA). After an equilibrium time of 10 min in ZEEP (step 2), PEEP was increased to 10 cm H₂O for 10 min (step 3). Thereafter, RM was performed, consisting of the application of continuous positive airway pressure (40 cm H₂O/40 s). In the event of a drop in systolic blood pressure by more than 20%, RM would have been terminated.

To analyze the specific effects of RM, 10 patients in each group were subjected to PEEP of 10 cm H₂O alone. For the others, immediately after RM was applied, PEEP was adjusted to 10 cm H₂O. In all patients, a second set of measurements was obtained at 5 min (step 4) and 30 min (step 5) postintervention (RM or no RM). A final measurement was performed after the pneumoperitoneum was released on completion of surgery. Apart from RM and the described changes in PEEP levels, basal ventilatory settings were kept constant for each patient throughout the experiment.

Physiologic Measurements

In both groups, preoperative lung function tests and EELV measurements were obtained using the helium dilution method (Spirodyne[®]; Dyn[®], Muret, France), in a 30° head-up position at end-expiration, to obtain reference EELV values when awake. Patients were asked to breathe normally (*i.e.*, at their usual tidal volume).

At each step of the experiment (before and after induction of pneumoperitoneum in ZEEP, PEEP 10 cm H₂O before intervention, 5 min and 30 min after intervention, and PEEP

10 cm H₂O after the pneumoperitoneum was exsufflated), peak ($P_{aw,max}$) and plateau end-inspiratory (P_{plat}) airway pressures were recorded using the end-inspiratory and end-expiratory occlusion technique.²³ Intrinsic PEEP was evaluated by means of end-expiratory occlusion. The quasistatic elastance of the total respiratory system (E_{rs}) was calculated as $\Delta P_{aw}/V_T$, where ΔP_{aw} is the difference between plateau end-inspiratory and end-expiratory airway pressure at a period of no-flow (corrected for intrinsic PEEP), and V_T is the tidal volume. EELV was measured twice (wash-out/wash-in method) using an automated procedure available on the ventilator (COVX module; GE Healthcare, Helsinki, Finland). EELV measurements, which reflect the amount of gas in the lungs,³⁹ require an inspired oxygen fraction step change of 0.1, without interruption of mechanical ventilation or any need for supplementary tracer gases, as previously described.^{40–42} Previous studies have specifically evaluated the reproducibility,⁴³ accuracy, and precision^{33,43,44} of EELV measurements provided by the ventilator.

To avoid any influence of the step change in FiO₂, arterial blood samples were taken from the radial artery before preoxygenation and at each step of the protocol just before EELV measurement. Arterial partial pressure of oxygen (PaO₂), arterial partial pressure of carbon dioxide (PaCO₂), and arterial pH were measured using a blood-gas analyzer (IL Synthesis; Instrumentation Laboratory[®], Lexington, MA).

The physiologic dead space to V_T ratio (V_D/V_T) was computed according to the following formula: $V_D/V_T = (PaCO_2 - PECO_2)/PaCO_2$, where PECO₂ is the mixed expired carbon dioxide partial pressure. Data were obtained by means of continuous expiratory air sampling, using a mainstream sensor placed in-line between the endotracheal tube and the Y-piece (CO₂ SMO PLUS 8100; Novamatrix Medical Sys-

Table 1. Patients Characteristics (N = 60)

Variable	Healthy Weight Study Group (n = 30)		Obese Study Group (n = 30)	
	PEEP + RM (n = 20)	PEEP (n = 10)	PEEP + RM (n = 20)	PEEP (n = 10)
Age, yr	48 ± 11	49 ± 7	46 ± 11	47 ± 5
Sex, m/f, no.	10/10	6/4	9/11	3/7
Weight, kg	65 ± 13	66 ± 11	140 ± 18*	132 ± 21*
Height, cm	170 ± 8	172 ± 7	169 ± 9	169 ± 7
Ideal body weight, kg	58 ± 11	60 ± 10	60 ± 10	62 ± 11
BMI, kg/m ²	22 ± 3	23 ± 1	46 ± 9*	45 ± 5*
ASA Classification, 1/2/3, no.	3/11/6	2/5/3	2/15/3	3/3/4

All data are presented as mean ± SD unless otherwise specified.

* $P < 0.05$ vs. healthy weight group.

ASA Classification = American Society of Anesthesiologists Physical Status Classification; BMI = body mass index; f = female; m = male; PEEP = 10 cm H₂O positive end-expiratory pressure; RM = recruitment maneuver.

tem Inc., Wallingford, CT). At each step of the procedure, the arterial-to-end-tidal partial pressure of CO₂ difference (Pa-ETCO₂) was also recorded. Pa-ETCO₂ has been found to closely correlate with atelectatic lung area on computed tomography (CT),⁴⁵ and is a valuable indicator of atelectasis during anesthesia and pneumoperitoneum.²⁷

Statistical Analysis

All test were performed using SEM software (version 2.0; center Jean Perrin, Clermont-Ferrand, France, ©1999).⁴⁶ Results are expressed as means ± SD. Based on previous data from our group and others,^{7,33,34} we calculated that at least 20 patients would be required in each group to allow a minimum detectable difference in mean EELV of at least 20% with an expected SD of 20%, assuming an α risk of 0.05 and a power of 0.9. In addition, to better compare the effects of RM when combined with PEEP and with PEEP alone, 10 more patients were included for each group.

A chi-square test was used to compare categorical data. Categorical and quantitative data were correlated using a Student *t* test or analysis of variance when normally distributed (and variances were equivalent), and the Kruskal-Wallis H-test otherwise. A two-way analysis of variance with different size groups was performed to test the effect of categorical parameters on longitudinal data. Comparisons of EELV, gas exchange, and respiratory mechanics between two points in time were performed using a paired Student *t* test. Inter-group differences were determined using the unpaired Student *t* test or the Kruskal-Wallis H-test. Within-group effect of PEEP with and without RM was analyzed using analysis of variance or the Kruskal-Wallis H-test, as appropriate. *Post hoc* analyses for pairwise comparisons were performed with the Bonferroni test. When applicable, correlations between variables were analyzed using Pearson coefficient correlation when the variables were normally distributed and Spearman ρ coefficient otherwise. Statistical testing was two-tailed with significance assumed at $P < 0.05$.

Results

The 60 patients approached for consent to participate in this study accepted. Data from all 60 patients are included in the analysis. Patient baseline characteristics are summarized in table 1. Except for weight and body mass index, baseline characteristics were similar between the study groups. Surgical procedures performed were laparoscopic gastric resection (n = 16), splenectomy (n = 8), and hepatectomy (n = 6) in the healthy weight group; laparoscopic sleeve gastrectomy (n = 18) and Roux-en-Y gastric bypass (n = 12) in the obese group. No relevant clinical problems occurred during any procedure or during surgery. No patients needed mechanical ventilation after the operation. Preoperative reference values of EELV were 2860 ml in healthy weight patients and 2170 ml in obese patients ($P < 0.01$).

During PEEP changes or RM, no significant differences in hemodynamic data were observed (tables 2 and 3).

Effects of Anesthesia Induction and Pneumoperitoneum Insufflation

There was no difference in the duration of anesthesia induction between the two groups (166 ± 13 vs. 159 ± 14 s, in the healthy weight and obese group, respectively; $P = 0.13$). Although no healthy weight patients developed hypoxic apnea, one obese patient did ($P = 0.31$). Compared with pre-induction values, anesthesia induction and mechanical ventilation with ZEEP significantly reduced PaO₂/FIO₂ ratios (healthy weight, 448 ± 72 vs. 341 ± 90 mmHg, $P < 0.01$; obese, 394 ± 75 mmHg vs. 214 ± 90 mmHg, $P < 0.01$) and EELV (−40 and −59%, in healthy weight and obese patients, respectively; both P values less than 0.01 vs. reference EELV values when awake).

Pneumoperitoneum further reduced EELV in the two groups (table 4, both P values less than 0.01 vs. EELV before pneumoperitoneum). Pneumoperitoneum increased PaCO₂ in the two study groups, whereas oxygenation was unaffected (tables 2 and 3). Pneumoperitoneum also increased Pa-

Table 2. Respiratory Mechanics, Gas Exchange, and Hemodynamic Data in Healthy Weight Patients (N = 30)

	ZEEP after Pneumoperitoneum	ZEEP before Pneumoperitoneum	PEEP 10 cm H ₂ O	Postintervention		End of Surgery
				5 min	30 min	
Peak airway pressure, cm H ₂ O	—	—	—	—	—	—
PEEP	17 ± 4	22 ± 3*	30 ± 2†	30 ± 4	30 ± 3	25 ± 4
PEEP + RM	15 ± 3	21 ± 4*	28 ± 3†	26 ± 3†‡	27 ± 3†	22 ± 2
End-inspiratory plateau pressure, cm H ₂ O	—	—	—	—	—	—
PEEP	12 ± 2	15 ± 2*	23 ± 5†	22 ± 3†	23 ± 2†	18 ± 4†
PEEP + RM	10 ± 3	14 ± 3*	21 ± 3†	18 ± 2†‡	19 ± 2†	16 ± 5†‡
Intrinsic PEEP, cm H ₂ O	—	—	—	—	—	—
PEEP	0.1 ± 0.2	0.2 ± 0.2	0.3 ± 0.2	0.3 ± 0.4	0.3 ± 0.5	0.1 ± 0.2
PEEP + RM	0.2 ± 0.6	0.2 ± 0.3	0.3 ± 0.4	0.4 ± 0.6	0.5 ± 0.5	0
E _{r,s} , cm H ₂ O/ml	—	—	—	—	—	—
PEEP	22 ± 4	30 ± 6*	25 ± 4†	25 ± 5†	23 ± 4	15 ± 3†
PEEP + RM	20 ± 7	31 ± 8*	24 ± 6†	18 ± 4†‡§	21 ± 6†	13 ± 4†‡
Arterial pH	—	—	—	—	—	—
PEEP	7.43 ± 0.02	7.42 ± 0.02	7.40 ± 0.03†	7.40 ± 0.02†	7.41 ± 0.02	7.42 ± 0.02
PEEP + RM	7.43 ± 0.02	7.44 ± 0.03	7.40 ± 0.02†	7.40 ± 0.03†	7.38 ± 0.05†	7.40 ± 0.04†
PaO ₂ , mmHg	—	—	—	—	—	—
PEEP	165 ± 52	152 ± 37	158 ± 33	156 ± 40	151 ± 37	160 ± 24
PEEP + RM	170 ± 49	173 ± 51	176 ± 46	207 ± 47†‡§	201 ± 56†‡§	210 ± 48†‡§
Paco ₂ , mmHg	—	—	—	—	—	—
PEEP	37 ± 3	40 ± 3*	37 ± 4†	37 ± 2†	39 ± 3	38 ± 5
PEEP + RM	37 ± 4	39 ± 2*	37 ± 2†	35 ± 2†‡§	38 ± 1	37 ± 3†
Pa-ETco ₂ , mmHg	—	—	—	—	—	—
PEEP	6 ± 3	9 ± 4*	6 ± 2†	6 ± 3†	7 ± 4	6 ± 3
PEEP + RM	6 ± 2	9 ± 3*	6 ± 1†	4 ± 1†‡§	5 ± 1†‡	4 ± 1†‡§
V _D /V _T	—	—	—	—	—	—
PEEP	0.15 ± 0.03	0.22 ± 0.04*	0.20 ± 0.03	0.20 ± 0.02	0.24 ± 0.03	0.18 ± 0.02†
PEEP + RM	0.12 ± 0.04	0.2 ± 0.06*	0.16 ± 0.04	0.11 ± 0.04†‡§	0.12 ± 0.03†‡	0.11 ± 0.03†‡
Mean arterial pressure, mmHg	—	—	—	—	—	—
PEEP	80 ± 4	82 ± 7	84 ± 5	83 ± 4	79 ± 5	77 ± 4
PEEP + RM	76 ± 6	78 ± 9	79 ± 7	78 ± 4	80 ± 4	77 ± 6
Heart rate, beats/min	—	—	—	—	—	—
PEEP	79 ± 12	79 ± 15	80 ± 23	80 ± 18	87 ± 9	83 ± 6
PEEP + RM	82 ± 15	80 ± 20	80 ± 14	84 ± 15	85 ± 10	80 ± 16

For all variables measured in healthy weight patients (N = 30), n = 10 for positive end-expiratory pressure (PEEP); n = 20, PEEP + recruitment maneuver (RM). All data are presented as mean ± SD unless otherwise specified.

* P < 0.01, vs. pneumoperitoneum after ZEEP. † P < 0.01, vs. pneumoperitoneum before ZEEP. ‡ P < 0.01, vs. PEEP 10 cm H₂O. § P < 0.01, PEEP + RM vs. PEEP.

E_{r,s} = elastance of the respiratory system; Paco₂ = arterial partial pressure of carbon dioxide; PaO₂ = arterial partial pressure of oxygen; Pa-ETco₂ = difference between arterial and end-tidal partial pressure of carbon dioxide; V_D/V_T = physiological dead space; ZEEP = zero end-expiratory pressure.

ETCO₂ and the V_D/V_T ratio (both P values less than 0.01). Overall respiratory mechanics worsened after pneumoperitoneum was induced (tables 2 and 3).

No intrinsic PEEP was detected in healthy weight or obese patients.

Effects of PEEP

Compared with ZEEP after pneumoperitoneum was induced, PEEP 10 cm H₂O significantly increased EELV in both study groups (healthy weight, 570 ml, P < 0.001; obese, 364 ml, P < 0.001). There was a significant difference between the two study groups (P = 0.003). In all patients, PEEP-induced changes for EELV were 46% (P < 0.001, compared with ZEEP after pneumoperitoneum) with no significant difference between PEEP+RM and PEEP alone

(table 4). PEEP also lowered Paco₂ and Pa-ETCO₂ in the two groups with no significant effect on oxygenation (tables 2 and 3). Overall respiratory mechanics improved after application of PEEP.

Effects of the Recruitment Maneuver

After RM, EELV further increased in both study groups (healthy weight, 154 ml; obese, 233 ml, fig. 2). Compared to PEEP alone, RM-induced changes of EELV were 10% in healthy weight and 20% in obese patients, with a statistically significant difference between the two groups (P = 0.026). In all patients, RM-induced change in EELV was 16% (P = 0.04). After RM, gas exchange also improved in the two study groups (tables 2 and 3). In contrast, PEEP alone did not cause any significant change in oxygenation. Compared

Table 3. Respiratory Mechanics, Gas Exchange, and Hemodynamic Data in Obese Patients (N = 30)

Variable	ZEEP after Pneumoperitoneum	ZEEP before Pneumoperitoneum	PEEP 10 cm H ₂ O	Postintervention		End of Surgery
				5 min	30 min	
Peak airway pressure, cm H ₂ O	—	—	—	—	—	—
PEEP	28 ± 5	34 ± 5*	37 ± 3†	37 ± 2	36 ± 4	33 ± 5
PEEP + RM	26 ± 4	31 ± 5*	33 ± 4†	31 ± 4†§	31 ± 5†§	30 ± 5‡
End-inspiratory plateau pressure, cm H ₂ O	—	—	—	—	—	—
PEEP	17 ± 3	20 ± 3*	23 ± 3†	23 ± 4†	24 ± 5†	22 ± 4
PEEP + RM	16 ± 3	19 ± 3*	22 ± 3†	19 ± 2†§	20 ± 3§	19 ± 2‡
Intrinsic PEEP, cm H ₂ O	—	—	—	—	—	—
PEEP	0.5 ± 0.5	0.5 ± 0.3	0.4 ± 0.6	0.4 ± 0.5	0.5 ± 0.3	0.5 ± 0.2
PEEP + RM	0.6 ± 1	0.5 ± 0.4	0.5 ± 0.5	0.7 ± 0.4	0.4 ± 0.5	0.4 ± 0.6
E _{rs} , cm H ₂ O/ml	—	—	—	—	—	—
PEEP	34 ± 4	40 ± 6*	28 ± 8†	28 ± 6	30 ± 7	27 ± 5†
PEEP + RM	31 ± 6	39 ± 10*	25 ± 7†	20 ± 5†‡§	22 ± 6†§	18 ± 5†‡§
Arterial pH	—	—	—	—	—	—
PEEP	7.42 ± 0.04	7.41 ± 0.03	7.39 ± 0.02†	7.39 ± 0.04†	7.38 ± 0.02†	7.39 ± 0.03†
PEEP + RM	7.44 ± 0.03	7.40 ± 0.02	7.38 ± 0.03†	7.39 ± 0.03†	7.37 ± 0.03†	7.38 ± 0.04†
PaO ₂ , mmHg	—	—	—	—	—	—
PEEP	102 ± 39	94 ± 28	104 ± 28	104 ± 20	98 ± 21	106 ± 15
PEEP + RM	107 ± 46	105 ± 43	111 ± 40	146 ± 37†‡§	149 ± 40†‡§	169 ± 42†‡§
Paco ₂ , mmHg	—	—	—	—	—	—
PEEP	42 ± 4	45 ± 5*	40 ± 5†	40 ± 4†	42 ± 3	41 ± 4
PEEP + RM	40 ± 3	44 ± 2*	41 ± 2†	39 ± 3†‡	41 ± 2	40 ± 3
Pa-ETCO ₂ , mmHg	—	—	—	—	—	—
PEEP	9 ± 1	14 ± 3*	10 ± 4†	10 ± 3†	11 ± 2†	9 ± 3†
PEEP + RM	10 ± 2	14 ± 2*	9 ± 3†	5 ± 1†‡§	6 ± 2†‡§	5 ± 2†‡
V _D /V _T	—	—	—	—	—	—
PEEP	0.30 ± 0.02	0.47 ± 0.3*	0.43 ± 0.03	0.42 ± 0.04	0.44 ± 0.02	0.39 ± 0.03
PEEP + RM	0.35 ± 0.04	0.45 ± 0.6*	0.40 ± 0.06	0.24 ± 0.06†‡§	0.32 ± 0.05†‡§	0.30 ± 0.05†‡
Mean arterial pressure, mmHg	—	—	—	—	—	—
PEEP	83 ± 24	87 ± 24	82 ± 10	82 ± 14	86 ± 10	80 ± 9
PEEP + RM	88 ± 17	85 ± 15	80 ± 20	82 ± 16	82 ± 10	84 ± 12
Heart rate, beats/min	—	—	—	—	—	—
PEEP	70 ± 15	72 ± 25	73 ± 12	74 ± 24	77 ± 15	79 ± 20
PEEP + RM	75 ± 16	74 ± 15	77 ± 18	80 ± 15	75 ± 14	74 ± 12

For all variables measured in obese patients (N = 30), n = 10 for positive end-expiratory pressure (PEEP); n = 20, PEEP + recruitment maneuver (RM). All data are presented as mean ± SD unless otherwise specified.

* $P < 0.01$, vs. pneumoperitoneum after ZEEP. † $P < 0.01$, vs. pneumoperitoneum before ZEEP. ‡ $P < 0.01$, vs. PEEP 10 cm H₂O. § $P < 0.05$, PEEP + RM vs. PEEP.

E_{rs} = elastance of the respiratory system; Paco₂ = arterial partial pressure of carbon dioxide; Pa-ETCO₂ = difference between arterial and end-tidal partial pressure of carbon dioxide; V_D/V_T = physiological dead space; ZEEP = zero end-expiratory pressure.

to PEEP alone, overall respiratory mechanics further improved after RM was performed (tables 2 and 3).

Thirty minutes after intervention, EELV was lower but still significant in the two groups compared to ZEEP after pneumoperitoneum was induced (table 4). There was no significant difference compared to PEEP 10 cm H₂O preintervention. The difference between the PEEP only and the PEEP+RM groups was significant only in healthy weight patients. Furthermore, in contrast to PEEP alone, 30 min after intervention, improvements in PaO₂, Pa-ETCO₂, and the V_D/V_T ratio were all sustained with PEEP+RM (tables 2 and 3).

End-expiratory Lung Volume, Static Elastance, Oxygenation, and Dead Space

A significant correlation was found between changes in EELV with pneumoperitoneum and changes in respiratory

system elastance ($r = 0.49$, $P < 0.01$) and Pa-ETCO₂ ($r = 0.39$, $P = 0.012$), as well as between changes in elastance and dead space after pneumoperitoneum ($r = 0.46$, $P < 0.01$). No correlation was found between change of EELV with pneumoperitoneum and change of oxygenation ($P = 0.18$).

A significant but weak correlation was found between PEEP-induced change in EELV (calculated using ZEEP during pneumoperitoneum as the reference) and change in respiratory system elastance ($r = 0.33$, $P = 0.03$) whereas no correlation was found between change in EELV with application of PEEP and oxygenation ($P = 0.90$). RM-induced changes in EELV correlated with changes in respiratory system elastance ($r = 0.51$, $P < 0.001$) and oxygenation ($r = 0.34$, $P = 0.03$). A significant correlation was also found between RM-induced change of EELV (with PEEP 10 cm H₂O before RM as the reference) and change of oxygenation ($r = 0.42$, $P < 0.01$).

Table 4. End-expiratory Lung Volume, ml (N = 60)

Measure	Healthy Weight Study Group (n = 30)		Obese Study Group (n = 30)	
	PEEP + RM (n = 20)	PEEP (n = 10)	PEEP + RM (n = 20)	PEEP (n = 10)
Pneumoperitoneum	—	—	—	—
After ZEEP	1,724 ± 774	1,802 ± 511	886 ± 284	934 ± 104
Before ZEEP	1,194 ± 405	1,225 ± 266	750 ± 258	771 ± 116
PEEP	1,750 ± 472*	1,680 ± 275*	1,115 ± 340*	1,017 ± 190*
Postintervention	—	—	—	—
5 min	1,958 ± 461*†‡	1,664 ± 341*	1,348 ± 317*†‡	1,013 ± 105*
30 min	1,928 ± 546*‡	1,541 ± 248*	1,277 ± 342*	983 ± 108

All data are presented as mean ± SD unless otherwise specified.

* $P < 0.001$ vs. pneumoperitoneum before ZEEP. † $P < 0.001$ vs. PEEP 10 cm H₂O. ‡ $P < 0.001$ PEEP + RM vs. PEEP only.

PEEP = positive end-expiratory pressure (10 cm H₂O); RM = recruitment maneuver; ZEEP = zero end-expiratory pressure.

Discussion

During pneumoperitoneum, in the beach chair position, 10 cm H₂O of PEEP improved EELV and respiratory elastance with no major change in oxygenation, whereas RM further improved EELV, respiratory elastance, and oxygenation in healthy weight and obese patients.

Pneumoperitoneum worsens respiratory mechanics in healthy weight^{4–6} and obese patients.^{7–10} Such effects may be related to reduced lung volumes and atelectasis formation, as previously confirmed by CT.¹² After induction of anesthesia, we found an average EELV of approximately 900 ml in obese patients and 1,700 ml in healthy weight patients, a result that is in line with measurements obtained by spiral CT and helium dilution technique in the absence of PEEP.^{16,23,47} Likewise, these results are in agreement with previous findings that pneumoperitoneum further reduced EELV after anesthesia induction in healthy weight^{4,47} and obese patients.^{7,48} Although morbid obesity *per se* may cause significant changes in respiratory system function and oxygenation,⁴⁹ the effects of pneumoperitoneum on oxygenation have been described as variable. Valenza *et al.*⁷ found that, despite severe impairment in respiratory mechanics and reduced EELV, oxygenation was improved during pneumoperitoneum. We confirm data indicating that oxygenation was not affected by pneumoperitoneum.^{6,17} Similar results were obtained in healthy weight patients. This finding is partly in agreement with the results of Sprung *et al.*,⁶ who reported that alterations in respiratory mechanics induced by pneumoperitoneum in healthy weight patients were greater than those seen in obese patients.

Ventilation at low EELV may cause or worsen lung injury, most likely as a result of the opening and closing of atelectatic lung regions and peripheral airways, whereas PEEP attenuates such injury.³⁵ Both PEEP and positioning (*e.g.*, reverse Trendelenburg position) have been found to attenuate the effects of pneumoperitoneum on respiratory mechanics and lung volumes.^{7,18,19} Among patients with a pneumoperitoneum-induced decrease in lung volume, PEEP increased EELV above the closing capacity.¹⁹ We

found that PEEP 10 cm H₂O only partly counteracted the detrimental effects of pneumoperitoneum on EELV and respiratory system elastance. No major effect on oxygenation was evident in either of our two study groups. Recent studies have also found that PEEP alone was insufficient to improve oxygenation during an increase in intraabdominal pressure.^{24,50} Indeed, PEEP may increase the normally aerated lung fraction in parallel with a reduction in the proportion of poorly aerated lung tissue although the extent of atelectasis may remain unchanged.²³

An RM has been proposed as valuable during pneumoperitoneum in obese patients.^{9,17,24} In contrast, few data are available on the use of RM in healthy weight patients. It has been shown in normal-weight patients that a single insufflation of 40 cm H₂O for 8 s was sufficient to open atelectatic areas after induction of anesthesia.⁵¹ In a recent study, Maisch *et al.*²⁸ demonstrated that, in normal-weight patients, RM in conjunction with PEEP 10 cm H₂O provided significant improvements in EELV, respiratory mechanics, and oxygenation during operation without any derangement of the lung and diaphragm position. We measured EELV, a sensitive indicator of airway collapse and PEEP-induced re-aeration,³⁶ and found that, compared to PEEP alone, the RM of 40 cm H₂O for 40 s was associated with average increases in EELV of 150 and 230 ml, respectively, for healthy weight and obese patients. We also observed a marked increase in oxygenation associated with improved respiratory system elastance, dead space, and Pa-ETCO₂. These observations are in agreement with previous results where RM decreased Pa-ETCO₂, a useful indicator of lung collapse and reopening after open-lung PEEP,⁴⁵ which in turn reduced dead space.^{45,52} Nevertheless, high insufflation pressures during RM may expose patients to hemodynamic instability, especially among those who are hypovolemic.⁵³ Therefore, we have taken special care that patients were normovolemic before performing RM. In addition, as an increase in EELV may result from alveolar recruitment and/or overdistension,^{33,39} we measured respiratory system elastance and dead space to distinguish between these possibili-

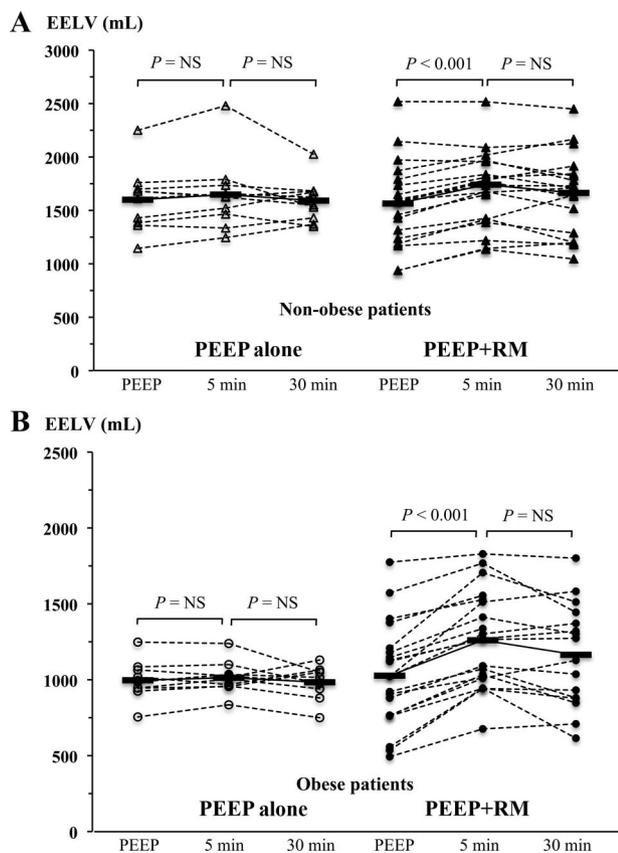


Fig. 2. Individual values of end-expiratory lung volume after pneumoperitoneum was insufflated in healthy weight (A) and obese (B) patients after the application of 10 cm H₂O positive end-expiratory pressure (PEEP) alone (hollow circle) or PEEP + recruitment maneuver (RM) (solid circle). End-expiratory lung volume (EELV) measurements were obtained at PEEP 10 cm H₂O as well as at 5 and 30 min after application of the recruitment maneuver (PEEP + RM) or with PEEP alone (PEEP). RM-induced changes of EELV were 10% in healthy weight patients and 20% in obese patients (both *P* values less than 0.001 vs. PEEP 10 cm H₂O), with a statistically difference (*P* = 0.026) between the two study groups. Bars represent mean values. NS = not (statistically) significant.

ties.^{28,30} However, we did not measure lung elastance, which may have been more relevant. During both pneumoperitoneum and beach chair position, PEEP-induced improvement in respiratory system elastance with PEEP was mainly related to changes in chest wall elastance.⁷ After pneumoperitoneum was induced, intraabdominal pressure and chest wall elastance both remained constant. Thus, it may be assumed that changes in elastance after RM mostly affected the lungs.

Although PEEP 10 cm H₂O was beneficial during pneumoperitoneum in healthy weight¹⁸ and obese patients,^{18,23,28} it may be argued that, when searching for safer ventilation, applying lower levels of PEEP to an open lung could be effective in healthy weight patients. It has been shown that application of PEEP 5 cm H₂O alone was effective in reducing atelectasis during pneumoperitoneum and improving oxygenation in normal-weight patients.⁵⁴ Our results are in contrast to previous findings for normal-weight patients where it was reported that

RM followed by ZEEP significantly reduced atelectasis for at least 20 min when using reduced FIO₂.⁵⁵ In addition, we found that, in contrast to PEEP alone, most healthy weight patients still improved in oxygenation and EELV after RM was performed without evidence for overdistension with PEEP 10 cm H₂O. Our results are consistent with those of Maisch *et al.*,²⁸ who reported that, in normal-weight patients without further EELV reduction by pneumoperitoneum insufflation, RM followed by high levels of PEEP is required to increase EELV effectively with reduced dead space.

Our study had several limitations. First, we did not measure the real extent of alveolar recruitment after application of PEEP and RM. Nevertheless, EELV measurements using the modified nitrogen wash-out/wash-in method correlate well with EELV as measured by CT,⁴¹ as well as with changes in lung aeration and consolidation gathered through CT.³⁶ Moreover, we excluded patients with airway disease (especially those with chronic obstructive pulmonary disease) for whom differences between CT measurement and ventilation-based assessment techniques may be relevant. Second, we did not conduct a detailed evaluation of hemodynamics during our investigation. However, a recent study⁵³ confirmed the hemodynamic safety of RM and application of PEEP in intravascular volume-loaded patients. Third, our procedural steps were not randomized. Owing to the specific procedures required by our study protocol and difficulties in collecting these measurements, it was difficult to design a randomized study. Fourth, the rather short interval between procedural steps is an additional limitation. However, the equilibration time of 10 min allowed readings to be within the accuracy limits issued by the instrument manufacturers. Moreover, as the sequence was the same in both groups, we believe our comparisons are valid. Fifth, although oxygenation and respiratory mechanics were sustainably improved at 30 min, we cannot exclude later variations in these parameters. Indeed, repeated RMs have been shown to improve both compliance and oxygenation, compared with a single RM.²⁴ Therefore, repeated RMs in conjunction with PEEP may represent an “optimal” open-lung approach.

Conclusion

Pneumoperitoneum worsens reduction in EELV and respiratory mechanics produced by anesthesia induction among normal-weight and obese patients, with no major effect on oxygenation. In contrast to PEEP alone, a PEEP of 10 cm H₂O combined with RM induces sustained improvements in EELV, gas exchange, and respiratory mechanics, and may be useful in counteracting the detrimental effects of pneumoperitoneum—especially on lung volume reduction in healthy weight and obese patients.

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References

1. Marshall BE, Wyche MQ, Jr: Hypoxemia during and after anesthesia. *ANESTHESIOLOGY* 1972; 37:178–209

2. Gunnarsson L, Strandberg A, Brismar B, Tokics L, Lundquist H, Hedenstierna G: Atelectasis and gas exchange impairment during enflurane/nitrous oxide anaesthesia. *Acta Anaesthesiol Scand* 1989; 33:629-37
3. Brismar B, Hedenstierna G, Lundquist H, Strandberg A, Svensson L, Tokics L: Pulmonary densities during anaesthesia with muscular relaxation—a proposal of atelectasis. *ANESTHESIOLOGY* 1985; 62:422-8
4. Pelosi P, Foti G, Cereda M, Vicardi P, Gattinoni L: Effects of carbon dioxide insufflation for laparoscopic cholecystectomy on the respiratory system. *Anaesthesia* 1996; 51: 744-9
5. Drummond GB, Martin LV: Pressure-volume relationships in the lung during laparoscopy. *Br J Anaesth* 1978; 50: 261-70
6. Sprung J, Whalley DG, Falcone T, Warner DO, Hubmayr RD, Hammel J: The impact of morbid obesity, pneumoperitoneum, and posture on respiratory system mechanics and oxygenation during laparoscopy. *Anesth Analg* 2002; 94:1345-50
7. Valenza F, Vagginielli F, Tiby A, Francesconi S, Ronzoni G, Guglielmi M, Zappa M, Lattuada E, Gattinoni L: Effects of the beach chair position, positive end-expiratory pressure, and pneumoperitoneum on respiratory function in morbidly obese patients during anaesthesia and paralysis. *ANESTHESIOLOGY* 2007; 107:725-32
8. Sprung J, Whalley DG, Falcone T, Wilks W, Navratil JE, Bourke DL: The effects of tidal volume and respiratory rate on oxygenation and respiratory mechanics during laparoscopy in morbidly obese patients. *Anesth Analg* 2003; 97: 268-74
9. Talab HF, Zabani IA, Abdelrahman HS, Bukhari WL, Mamoun I, Ashour MA, Sadeq BB, El Sayed SI: Intraoperative ventilatory strategies for prevention of pulmonary atelectasis in obese patients undergoing laparoscopic bariatric surgery. *Anesth Analg* 2009; 109:1511-6
10. Dumont L, Mattys M, Mardirosoff C, Vervloesem N, Allé JL, Massaut J: Changes in pulmonary mechanics during laparoscopic gastropasty in morbidly obese patients. *Acta Anaesthesiol Scand* 1997; 41:408-13
11. Perilli V, Sollazzi L, Bozza P, Modesti C, Chierichini A, Tacchino RM, Ranieri R: The effects of the reverse trendelenburg position on respiratory mechanics and blood gases in morbidly obese patients during bariatric surgery. *Anesth Analg* 2000; 91:1520-5
12. Andersson LE, Bååth M, Thörne A, Aspelin P, Odeberg-Wernerman S: Effect of carbon dioxide pneumoperitoneum on development of atelectasis during anaesthesia, examined by spiral computed tomography. *ANESTHESIOLOGY* 2005; 102:293-9
13. El-Dawlatly AA, Al-Dohayan A, Abdel-Meguid ME, El-Bakry A, Manaa EM: The effects of pneumoperitoneum on respiratory mechanics during general anaesthesia for bariatric surgery. *Obes Surg* 2004; 14:212-5
14. Nguyen NT, Wolfe BM: The physiologic effects of pneumoperitoneum in the morbidly obese. *Ann Surg* 2005; 241:219-26
15. Pelosi P, Rocco PR: Airway closure: The silent killer of peripheral airways. *Crit Care* 2007; 11:114
16. Pelosi P, Ravagnan I, Giurati G, Panigada M, Bottino N, Tredici S, Eccher G, Gattinoni L: Positive end-expiratory pressure improves respiratory function in obese but not in normal subjects during anaesthesia and paralysis. *ANESTHESIOLOGY* 1999; 91:1221-31
17. Whalen FX, Gajic O, Thompson GB, Kendrick ML, Que FL, Williams BA, Joyner MJ, Hubmayr RD, Warner DO, Sprung J: The effects of the alveolar recruitment maneuver and positive end-expiratory pressure on arterial oxygenation during laparoscopic bariatric surgery. *Anesth Analg* 2006; 102:298-305
18. Maracajá-Neto LF, Verçosa N, Roncally AC, Giannella A, Bozza FA, Lessa MA: Beneficial effects of high positive end-expiratory pressure in lung respiratory mechanics during laparoscopic surgery. *Acta Anaesthesiol Scand* 2009; 53:210-7
19. Meininger D, Byhahn C, Mierdl S, Westphal K, Zwissler B: Positive end-expiratory pressure improves arterial oxygenation during prolonged pneumoperitoneum. *Acta Anaesthesiol Scand* 2005; 49:778-83
20. Kim JY, Shin CS, Kim HS, Jung WS, Kwak HJ: Positive end-expiratory pressure in pressure-controlled ventilation improves ventilatory and oxygenation parameters during laparoscopic cholecystectomy. *Surg Endosc* 2010; 24:1099-103
21. Rothen HU, Sporre B, Engberg G, Wegenius G, Hedenstierna G: Re-expansion of atelectasis during general anaesthesia: A computed tomography study. *Br J Anaesth* 1993; 71:788-95
22. Tusman G, Böhm SH, Vazquez de Anda GF, do Campo JL, Lachmann B: 'Alveolar recruitment strategy' improves arterial oxygenation during general anaesthesia. *Br J Anaesth* 1999; 82:8-13
23. Reinius H, Jonsson L, Gustafsson S, Sundbom M, Duvernoy O, Pelosi P, Hedenstierna G, Fredén F: Prevention of atelectasis in morbidly obese patients during general anaesthesia and paralysis: A computerized tomography study. *ANESTHESIOLOGY* 2009; 111:979-87
24. Almarakbi WA, Fawzi HM, Alhashemi JA: Effects of four intraoperative ventilatory strategies on respiratory compliance and gas exchange during laparoscopic gastric banding in obese patients. *Br J Anaesth* 2009; 102:862-8
25. Chalhoub V, Yazigi A, Sleilaty G, Haddad F, Noun R, Madi-Jebara S, Yazbeck P: Effect of vital capacity manoeuvres on arterial oxygenation in morbidly obese patients undergoing open bariatric surgery. *Eur J Anaesthesiol* 2007; 24:283-8
26. Neumann P, Rothen HU, Berglund JE, Valtysson J, Magnusson A, Hedenstierna G: Positive end-expiratory pressure prevents atelectasis during general anaesthesia even in the presence of a high inspired oxygen concentration. *Acta Anaesthesiol Scand* 1999; 43:295-301
27. Strang CM, Hachenberg T, Fredén F, Hedenstierna G: Development of atelectasis and arterial to end-tidal PCO₂-difference in a porcine model of pneumoperitoneum. *Br J Anaesth* 2009; 103:298-303
28. Maisch S, Reissmann H, Fuellekrug B, Weismann D, Rutkowski T, Tusman G, Bohm SH: Compliance and dead space fraction indicate an optimal level of positive end-expiratory pressure after recruitment in anesthetized patients. *Anesth Analg* 2008; 106:175-81
29. Constantin JM, Cayot-Constantin S, Roszyk L, Futier E, Sapin V, Dastugue B, Bazin JE, Rouby JJ: Response to recruitment maneuver influences net alveolar fluid clearance in acute respiratory distress syndrome. *ANESTHESIOLOGY* 2007; 106:944-51
30. Suarez-Sipmann F, Böhm SH, Tusman G, Pesch T, Thamm O, Reissmann H, Reske A, Magnusson A, Hedenstierna G: Use of dynamic compliance for open lung positive end-expiratory pressure titration in an experimental study. *Crit Care Med* 2007; 35:214-21
31. Richard JC, Maggiore SM, Mercat A: Clinical review: Bedside assessment of alveolar recruitment. *Crit Care* 2004; 8:163-9
32. Hedenstierna G: The recording of FRC—is it of importance and can it be made simple? *Intensive Care Med* 1993; 19:365-6
33. Bikker IG, van Bommel J, Reis Miranda D, Bakker J, Gommers D: End-expiratory lung volume during mechanical ventilation: A comparison with reference values and the effect of positive end-expiratory pressure in intensive care unit patients with different lung conditions [abstract]. *Crit Care* 2008; 12:R145

34. Futier E, Constantin JM, Petit A, Jung B, Kwiatkowski F, Duclos M, Jaber S, Bazin JE: Positive end-expiratory pressure improves end-expiratory lung volume but not oxygenation after induction of anaesthesia. *Eur J Anaesthesiol* 2010; 27:508-13
35. Duggan M, Kavanagh BP: Pulmonary atelectasis: A pathogenic perioperative entity. *ANESTHESIOLOGY* 2005; 102:838-54
36. Rylander C, Högman M, Perchiazzi G, Magnusson A, Hedenstierna G: Functional residual capacity and respiratory mechanics as indicators of aeration and collapse in experimental lung injury. *Anesth Analg* 2004; 98:782-9
37. Gander S, Frascarolo P, Suter M, Spahn DR, Magnusson L: Positive end-expiratory pressure during induction of general anesthesia increases duration of nonhypoxic apnea in morbidly obese patients. *Anesth Analg* 2005; 100:580-4
38. Lopes MR, Oliveira MA, Pereira VO, Lemos IP, Auler JO, Jr., Michard F: Goal-directed fluid management based on pulse pressure variation monitoring during high-risk surgery: A pilot randomized controlled trial [abstract]. *Crit Care* 2007; 11:R100
39. Heinze H, Eichler W: Measurements of functional residual capacity during intensive care treatment: The technical aspects and its possible clinical applications. *Acta Anaesthesiol Scand* 2009; 53:1121-30
40. Olegård C, Söndergaard S, Hoults E, Lundin S, Stenqvist O: Estimation of functional residual capacity at the bedside using standard monitoring equipment: A modified nitrogen washout/washin technique requiring a small change of the inspired oxygen fraction. *Anesth Analg* 2005; 101:206-12
41. Chiumello D, Cressoni M, Chierichetti M, Tallarini F, Botticelli M, Berto V, Mietto C, Gattinoni L: Nitrogen washout/washin, helium dilution and computed tomography in the assessment of end expiratory lung volume [abstract]. *Crit Care* 2008; 12:R150
42. Lambermont B, Ghuysen A, Janssen N, Morimont P, Hartstein G, Gerard P, D'Orio V: Comparison of functional residual capacity and static compliance of the respiratory system during a positive end-expiratory pressure (PEEP) ramp procedure in an experimental model of acute respiratory distress syndrome [abstract]. *Crit Care* 2008; 12:R91
43. Bikker IG, Scohy TV, Ad J J C Bogers, Bakker J, Gommers D: Measurement of end-expiratory lung volume in intubated children without interruption of mechanical ventilation. *Intensive Care Med* 2009; 35:1749-53
44. Scohy TV, Bikker IG, Hofland J, de Jong PL, Bogers AJ, Gommers D: Alveolar recruitment strategy and PEEP improve oxygenation, dynamic compliance of respiratory system and end-expiratory lung volume in pediatric patients undergoing cardiac surgery for congenital heart disease. *Paediatr Anaesth* 2009; 19:1207-12
45. Tusman G, Suarez-Sipmann F, Böhm SH, Pech T, Reissmann H, Meschino G, Scandurra A, Hedenstierna G: Monitoring dead space during recruitment and PEEP titration in an experimental model. *Intensive Care Med* 2006; 32:1863-71
46. Kwiatkowski F, Girard M, Hacene K, Berlie J: [Sem: A suitable statistical software (adapted) for research in oncology.] *Bull Cancer* 2000; 87:715-21
47. Pelosi P, Croci M, Ravagnan I, Tredici S, Pedoto A, Lissoni A, Gattinoni L: The effects of body mass on lung volumes, respiratory mechanics, and gas exchange during general anesthesia. *Anesth Analg* 1998; 87:654-60
48. Damia G, Mascheroni D, Croci M, Tarenzi L: Perioperative changes in functional residual capacity in morbidly obese patients. *Br J Anaesth* 1988; 60:574-8
49. Pelosi P, Croci M, Ravagnan I, Cerisara M, Vicardi P, Lissoni A, Gattinoni L: Respiratory system mechanics in sedated, paralyzed, morbidly obese patients. *J Appl Physiol* 1997; 82:811-8
50. Tafer N, Nouette-Gaulain K, Richebé P, Rozé H, Lafargue M, Janvier G: [Effectiveness of a recruitment manoeuvre and positive end-expiratory pressure on respiratory mechanics during laparoscopic bariatric surgery.] *Ann Fr Anesth Reanim* 2009; 28:130-4
51. Rothen HU, Neumann P, Berglund JE, Valtysson J, Magnusson A, Hedenstierna G: Dynamics of re-expansion of atelectasis during general anaesthesia. *Br J Anaesth* 1999; 82:551-6
52. Fletcher R, Jonson B, Cumming G, Brew J: The concept of deadspace with special reference to the single breath test for carbon dioxide. *Br J Anaesth* 1981; 53:77-88
53. Bohm SH, Thamm OC, von Sandersleben A, Bangert K, Langwieler TE, Tusman G, Strate TG, Standl TG: Alveolar recruitment strategy and high positive end-expiratory pressure levels do not affect hemodynamics in morbidly obese intravascular volume-loaded patients. *Anesth Analg* 2009; 109:160-3
54. Kim JY, Shin CS, Kim HS, Jung WS, Kwak HJ: Positive end-expiratory pressure in pressure-controlled ventilation improves ventilatory and oxygenation parameters during laparoscopic cholecystectomy. *Surg Endosc* 2010; 24:1099-103
55. Rothen HU, Sporre B, Engberg G, Wegenius G, Högman M, Hedenstierna G: Influence of gas composition on recurrence of atelectasis after a reexpansion maneuver during general anesthesia. *ANESTHESIOLOGY* 1995; 82:832-42