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Spontaneous breathing trial and post-extubation work of breathing in morbidly obese critically ill patients

Martin Mahul1†, Boris Jung1,4†, Fabrice Galia1, Nicolas Molinari2, Audrey de Jong1, Yannaël Coisel1,4, Rosanna Vaschetto3, Stefan Matecki4, Gérald Chanques1,4, Laurent Brochard5,6 and Samir Jaber1,4*

Abstract

Background: Predicting whether an obese critically ill patient can be successfully extubated may be specially challenging. Several weaning tests have been described but no physiological study has evaluated the weaning test that would best reflect the post-extubation inspiratory effort.

Methods: This was a physiological randomized crossover study in a medical and surgical single-center Intensive Care Unit, in patients with body mass index (BMI) >35 kg/m² who were mechanically ventilated for more than 24 h and underwent a weaning test. After randomization, 17 patients were explored using five settings: pressure support ventilation (PSV) 7 and positive end-expiratory pressure (PEEP) 7 cmH2O; PSV 0 and PEEP 7 cmH2O; PSV 7 and PEEP 0 cmH2O; PSV 0 and PEEP 0 cmH2O; and a T piece, and after extubation. To further minimize interaction between each setting, a period of baseline ventilation was performed between each step of the study. We hypothesized that the post-extubation work of breathing (WOB) would be similar to the T-tube WOB.

Results: Respiratory variables and esophageal and gastric pressure were recorded. Inspiratory muscle effort was calculated as the esophageal and trans-diaphragmatic pressure time products and WOB. Sixteen obese patients (BMI 44 kg/m² ± 8) were included and successfully extubated. Post-extubation inspiratory effort, calculated by WOB, was 1.56 J/L ± 0.50, not statistically different from the T piece (1.57 J/L ± 0.56) or PSV 0 and PEEP 0 cmH2O (1.58 J/L ± 0.57), whatever the index of inspiratory effort. The three tests that maintained pressure support statistically underestimated post-extubation inspiratory effort (WOB 0.69 J/L ± 0.31, 1.15 J/L ± 0.39 and 1.09 J/L ± 0.49, respectively, p < 0.001). Respiratory mechanics and arterial blood gases did not differ between the five tests and the post-extubation condition.

Conclusions: In obese patients, inspiratory effort measured during weaning tests with either a T-piece or a PSV 0 and PEEP 0 was not different to post-extubation inspiratory effort. In contrast, weaning tests with positive pressure overestimated post-extubation inspiratory effort.

Trial registration: Clinical trial.gov (reference NCT01616901), 2012, June 4th

Keywords: Weaning, Mechanical ventilation, Obese, Work of breathing, Acute respiratory failure

* Correspondence: s-jaber@chu-montpellier.fr
† Equal contributors

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Background
Exubation is a critical decision in the Intensive Care Unit (ICU). Exubation failure may occur in up to 20% [1] of patients and is associated with morbidity. Excessive and non-sustainable work of breathing (WOB) is likely a major reason for exubation failure [2–5]. Evaluation of how the critically ill patient is breathing with no assistance or a minimal level of assistance (the period known as the weaning test or the spontaneous breathing trial) [4] is therefore recommended before exubation [3, 4, 6, 7]. Different weaning tests are suggested for non-selected adult patients: a T-piece trial (oxygen supply without positive pressure), continuous positive airway pressure (CPAP) and low pressure support ventilation (PSV), with a low level of PSV, from 5 to 8 cmH2O, to compensate for the imposed workload due to the ventilator circuit [3, 4, 6, 7]. Although these weaning tests are not equivalent in term of the WOB [8, 9] and studies are underpowered to assess the risk of exubation failure, they are recommended to assess whether a patient is ready to be exubated [3, 6].

Predicting whether an obese critically ill patient can be successfully exubated may be specially challenging. Obesity decreases respiratory system compliance, inspiratory and expiratory lung volumes, functional residual capacity, upper airway mechanical function and neuromuscular strength [10]. Moreover, in obese patients, oxygen consumption is increased, with a high proportion of this consumption spent in the WOB [11–13]. Although the T piece, CPAP and low PSV levels have been used to reproduce post-exubation conditions in non-selected critically ill patients, the weaning test modality that would best reproduce post-exubation inspiratory effort (WOB and pressure time product indexes) in obese critically ill patients has never been evaluated and many clinicians are worried about using no support during the test [14, 15].

The aim of our study was thus to assess which weaning test would best reproduce post-exubation inspiratory effort in obese critically ill patients. We compared a T-piece trial to weaning tests with PSV 7 and positive end-expiratory pressure (PEEP) 7 cmH2O; PSV 0 and PEEP 7 cmH2O; PSV 7 and PEEP 0 cmH2O; PSV 0 and PEEP 0 cmH2O, in this particular population. We hypothesized that the T-tube or PSV 0 and PEEP 0 cmH2O would best approximate the post-exubation WOB.

Methods
Study
This was a physiological prospective randomized cross-over study (Additional file 1: Table S1), approved by the Ethics Committee of the Saint-Eloi Teaching Hospital (2012 A-00294-39, Comité de Protection des Personnes Sud Méditerranée III, Montpellier, France), and registered on clinical trial.gov (reference NCT01616901, registered June 4th, 2012). All patients provided their written informed consent.

Patients
Upon admission, height and weight were measured using the bed scale and a tape measure. All morbidly obese patients, defined by a body mass index (total body weight in kg/height in m²) >35 kg/m² [16], were considered eligible for inclusion in the study if they were mechanically ventilated for at least 24 h and were considered by the physician on duty to be ready for exubation. Patients were not included in the study if there was any contraindication to the insertion of an esophageal catheter.

Experimental procedure and study design
A 15-minute period corresponding to a baseline state was first recorded (using PSV and PEEP set by the clinician in charge of the patient before inclusion). Patients were then randomly assessed using computer-driven software with five settings: PSV 7 and PEEP 7 cmH2O; PSV 0 and PEEP 7 cmH2O; PSV 7 and PEEP 0 cmH2O; PSV 0 and PEEP 0 cmH2O or the T piece. Each setting lasted 15 minutes with a 10-minute period of return to baseline steady state between each setting (Fig. 1). Steady state was defined clinically as a period sufficient to ensure clinical stability in respiratory and hemodynamic variables assessed by a physical exam which took into account heart rate, respiratory rate, paradoxical breathing pattern, accessory muscle use, grunting at end expiration and nasal flaring [17], and as previously performed by our group [18, 19].

After being explored with these five settings, and in the case of clinical success in the different weaning trials, patients were ventilated for 10 minutes using baseline state variables and then were exubated if the clinical state was judged adequate by the clinician in charge. A post-exubation measurement was performed 20 minutes after exubation using an oro-nasal oxygen mask with a flow of 5 L/minute (equivalent to inspired oxygen fraction (FiO2) of 0.4 [20]). According to our local protocol described in detail in a previous review [21], and after having achieved each step of the protocol, non-invasive ventilation was performed as a prophylactic routine measure in the immediate post-exubation period, for between 30 and 45 minutes every 4 to 6 h. Settings were adjusted to target the following: tidal volume (VT) 6–10 ml/kg of ideal body weight, respiratory rate (RR) 12–20 c/minute and pulse...
arterial oxygen saturation (SpO2) equal or above 95%. Non-invasive ventilation was never performed before the end of the protocol.

Measurements
All patients were studied in a semi-recumbent position with the head of the bed elevated to an angle from 30 to 45 degrees, according to patient comfort. [22] Procedures are detailed in the additional material. Briefly, the respiratory mechanics measurements comprised flow, airway pressure, esophageal (Pes) and gastric (Pga) pressure swings. Trans-diaphragmatic swings (Pdi) were calculated by subtracting Pes from Pga. Minute ventilation (VE), tidal volume (Vt), inspiratory (Ti), expiratory time (Te), total cycle duration (Ttot) and RR were calculated from the numerical integration of the flow signal.

The inspiratory WOB per breath performed by the patient was calculated from a Campbell diagram taking into account the presence of intrinsic PEEP. Esophageal and trans-diaphragmatic pressure-time products (PTPes and PTPdi) were also measured as previously reported [23, 24]. Analyses of arterial blood gases were obtained at the end of each test.

Statistical analysis
All values are presented as mean ± SD. To assess differences between the weaning tests, we used the Friedman test and then pairwise comparisons with the Wilcoxon test if a significant difference appeared. Statistical analysis was performed by an independent statistician (NM) using R software® (R Foundation for Statistical Computing, Auckland, New Zealand).

Based on the literature review, we hypothesized that the post-extubation WOB would be similar to the T-tube WOB [25, 26] and would approximate 1.5 +/- 0.9 J/L in obese critically ill patients. We also hypothesized that WOB in PSV 7 cmH2O and PEEP 7 cmH2O would approximate 0.7 +/- 0.5 J/L [27]. Then, with an alpha risk at 0.05 and a power at 0.90, 12 patients would be needed. We decided to include 17 patients in order to make sure that 12 patients would complete the study. Significance was set at \( p < 0.01 \) after correction for the number of multiple comparisons, i.e., using the Bonferroni test.

Results
Patients
Between March and December 2012, 40 obese patients with body mass index ≥35 kg/m² were admitted in our center. Among them, 17 met the inclusion criteria. Sixteen patients (13 women and 3 men) with mean body mass index of 44 kg/m² (±8 kg/m²) were prospectively enrolled in the present study, as shown in Fig. 2. Characteristics of the subjects are detailed in Table 1. Mean duration of invasive mechanical
ventilation before enrollment in the study was 6 days (±7 days). The five weaning tests were well-tolerated by all patients and all of them but one were successfully extubated.

The first patient was initially unable to complete the five weaning tests. She was re-challenged 72 h later, and succeeded the tests and extubation. Seven days after extubation, she developed cardiorespiratory distress and was re-intubated. Patient number 15 developed hypoxemic acute respiratory failure and was re-intubated 12 h after extubation. One patient had accidental nasogastric catheter removal after extubation, preventing the measurement of respiratory muscle work variables after extubation. This patient was excluded from the final analysis.

**Respiratory variables and gas exchange**

There was no statistical difference in any of the different respiratory variables (shown in Table 2) among the five weaning tests or at 20 minutes after extubation. In particular, differences in the RR/V̇T ratio were not statistically significant between the five weaning
Table 1 Characteristics of the patients

| Patient number | Sex | Age (years) | SAPS II | Height (cm) | Weight (kg) | BMI (kg/m²) | Underlying diseases | Etiology of respiratory failure | ETT ID (mm) | MV before extubation (days) | PSV at baseline (cmH₂O) | PEEP at baseline (cmH₂O) | Extubation failure (Y/N) | Outcome (D/S) |
|----------------|-----|-------------|---------|-------------|-------------|-------------|---------------------|--------------------------------|-------------|-----------------------------|------------------------|------------------------|------------------------|----------------|---|
| 1              | F   | 83          | 150     | 80          | 35          | CHF         | Small bowel ischemia | 7.5                           | 7            | 8                          | 6                      | N                      | D                      |                |
| 2              | F   | 85          | 163     | 115         | 43          | NIDDM       | Pneumonia           | 7.5                           | 4            | 15                         | 7                      | N                      | S                      |                |
| 3              | M   | 64          | 170     | 130         | 44          | NIDDM       | Acute pancreatitis  | 8                             | 3            | 12                         | 8                      | N                      | S                      |                |
| 4              | F   | 59          | 155     | 95          | 39          | None        | Peritonitis         | 7.5                           | 3            | 12                         | 8                      | N                      | S                      |                |
| 5              | F   | 49          | 160     | 174         | 67          | COPD, OSA   | Septic shock       | 7.5                           | 6            | 10                         | 7                      | N                      | S                      |                |
| 6              | F   | 25          | 172     | 145         | 49          | None        | Asthma             | 7.5                           | 1            | 10                         | 8                      | N                      | S                      |                |
| 7              | F   | 54          | 153     | 121         | 51          | Asthma, HTN | Post abdominal surgery | 7.5                           | 1            | 10                         | 8                      | N                      | S                      |                |
| 8              | M   | 37          | 180     | 130         | 40          | None        | Acute pancreatitis  | 8                             | 14           | 8                          | 10                     | N                      | S                      |                |
| 9              | F   | 78          | 155     | 87          | 36          | None        | Bowel obstruction  | 7.5                           | 4            | 8                          | 5                      | N                      | S                      |                |
| 10             | F   | 49          | 167     | 112         | 41          | Asthma, OSA | Peritonitis         | 7.5                           | 30           | 8                          | 5                      | N                      | S                      |                |
| 11             | F   | 73          | 150     | 93          | 41          | CHF, AF     | Septic shock       | 7.5                           | 4            | 12                         | 6                      | N                      | D                      |                |
| 12             | F   | 50          | 162     | 94          | 36          | None        | Necrotizing fasciitis | 7.5                           | 2            | 9                          | 7                      | N                      | S                      |                |
| 13             | M   | 63          | 175     | 180         | 56          | NIDDM, HTN  | Small bowel bleeding | 7.5                           | 1            | 8                          | 10                     | N                      | S                      |                |
| 14             | F   | 43          | 155     | 105         | 43          | OSA, home ventilation | Pneumonia       | 7.5                           | 3            | 12                         | 7                      | N                      | S                      |                |
| 15             | F   | 77          | 155     | 84          | 36          | NIDDM, HTN  | Pancreatitis       | 7.5                           | 7            | 9                          | 7                      | Y                      | S                      |                |
| 16             | F   | 50          | 164     | 124         | 46          | OSA, home ventilation | Post abdominal surgery | 7.5                           | 8            | 14                         | 8                      | N                      | S                      |                |
| Mean           |     | 59          | 162     | 117         | 44          |             |                    |                               | 6            | 10                         | 7                      |                       |                       |                |
| SD             |     | 17          | 22      | 9           | 30          | 8           |             | 7                             | 2            | 2                          |                       |                       |                       |                |

Abbreviations: AF atrial fibrillation, BMI body mass index; CHF chronic heart failure, D deceased; ETT ID endotracheal tube internal diameter; F female; M male; HTN hypertension, mechanical ventilation; NIDDM non-insulin-dependent diabetes mellitus; OSA obstructive sleep apnoea, PEEP positive end-expiratory pressure; PSV pressure support ventilation; SAPS II Simplified Acute Physiology Score II [34]; S survived
There was no statistically significant difference in arterial blood gases or hemodynamic variables among the six steps of the study, as shown in Table 3.

**Table 3** Arterial blood gases and hemodynamic variables during the five different weaning tests and at 20 minutes after extubation

|                | PSV +7 cmH2O PEEP | PSV +7 cmH2O PEEP | PSV +7 cmH2O PEEP | PSV +7 cmH2O PEEP | T piece | After extubation |
|----------------|-------------------|-------------------|-------------------|-------------------|---------|------------------|
|                | +7 cmH2O PEEP     | 0 cmH2O PEEP      | +7 cmH2O PEEP     | 0 cmH2O PEEP      |         |                  |
| **Ph**, mmHg   | 7.45 ± 0.06       | 7.44 ± 0.06       | 7.44 ± 0.06       | 7.44 ± 0.06       | 7.43 ± 0.06 | 7.42 ± 0.06      |
| **PaCO2**, mmHg| 41 ± 11           | 42 ± 11           | 43 ± 12           | 43 ± 12           | 44 ± 13 | 44 ± 10          |
| **PaO2/FiO2**  | 277 ± 76          | 257 ± 81          | 252 ± 73          | 230 ± 65          | 217 ± 65 | 224 ± 51         |
| **SBP**, mmHg  | 148 ± 22          | 148 ± 26          | 148 ± 26          | 146 ± 30          | 150 ± 18 | 147 ± 24         |
| **DBP**, mmHg  | 72 ± 12           | 71 ± 12           | 73 ± 12           | 72 ± 15           | 69 ± 13 | 70 ± 15          |
| **HR**, beats/minute | 96 ± 14 | 97 ± 16 | 98 ± 16 | 100 ± 16 | 99 ± 14 | 101 ± 15 |

There were no statistically significant differences between respiratory variables among the successive tests. Abbreviations: DBP diastolic blood pressure, HR heart rate, ND not done, PEEP positive end-expiratory pressure, PSV pressure support ventilation, SBP systolic blood pressure.
An ongoing multicenter observational study in France (FREEREA study), will provide some epidemiological data about weaning and extubation in this particular population. The preliminary results (unpublished) show that among 64 critically ill morbidly obese patients extubated, 22 (34 %) were extubated after a T tube, 28 (44 %) after a low PSV trial, 12 (19 %) with no spontaneous breathing trial and 2 (3 %) after a different weaning trial. These data justify our study as there is wide heterogeneity of extubation practice in this population, with a high proportion of patients being extubated from a substantial level of support.

Our study presents limitations. First, we investigated the inspiratory effort indexes twenty minutes after extubation and the study was not designed to explore long-term consequences of several weaning tests on oxygenation, end-expiratory lung volume or outcome. Because outcome

Fig. 3 Esophageal (a) and trans-diaphragmatic (b) swings. Individual and mean changes in esophageal and trans-diaphragmatic swings during the five weaning tests and 20 minutes after extubation. All the tests show that the weaning tests that best reproduce respiratory muscle work after extubation were pressure support ventilation (PSV) 0 cmH₂O + positive end-expiratory pressure (PEEP) 0 cmH₂O and the T piece, with no statistically significant difference between the two. *p < 0.001 when compared with after extubation. Pdi transdiaphragmatic pressure, pes esophageal pressure.
was not a study endpoint, we cannot make any final recommendation about which weaning test is associated with the highest rate of weaning success. Ideally, a weaning test would perfectly predict the ability of the patient to breathe alone and without being ventilatory assisted by simulating the post-extubation respiratory constraint [26]. Second, post-extubation intermittent non-invasive ventilation is routinely used in our unit for high-risk patients [14, 21] to rest the inspiratory muscles and improve lung aeration. It may have contributed to our low rate of reintubation (6 %).

The present study focused on morbidly obese patients and found results consistent with the studies published by Straus et al. [25] and Cabello et al. [8], which included non-obese patients. We report that the T piece and PSV 0 + PEEP 0 cmH2O weaning tests were the two tests that best approximated the WOB after extubation. We also found that the PSV 7 + PEEP 0 cmH2O test leads to a major underestimation of the WOB after extubation in obese patients with significantly less inspiratory effort in comparison with both the T piece test and 20 minutes after extubation. Straus et al demonstrated...
that post-extubation WOB was well-approximated by the WOB during a T-piece test and that the endotracheal tube was responsible for about 11% of the total work of breathing. [25] More recently, Cabello et al. compared a spontaneous breathing trial on a T-piece with low PSV (7 cm H₂O) with or without PEEP in a subpopulation of patients with heart failure who were difficult to wean. [8] The authors concluded that performing the weaning test while maintaining a positive pressure in the circuit underestimates the postextubation WOB and unmasks a possible effect on left ventricular function, and suggested the T piece as the weaning test of choice in these patients.

In a landmark physiological study, Brochard et al. demonstrated that breathing through the T piece overestimates the WOB by 27 ± 18% compared to the postextubation period [26]. Contrary to the present study, Brochard et al. included a high proportion of patients...
with chronic obstructive pulmonary disease, and used ventilators with higher ventilatory circuit-resistive load [28] and lower pressurization performance, especially in terms of inspiratory-trigger-imposed WOB [29, 30].

As compared to the literature on non-obese patients, WOB values evaluated in the present study were higher [26, 31]. In morbidly obese patients, an elevation of pharyngeal collapsibility and upper airway resistance related to fatty deposits on pharynx and oral soft tissue and associated with local inflammation can increase the WOB [32]. Weaning trials performed with positive pressure underestimated post-extubation WOB by 33 % (0.5 J/L) up to 50 % (0.8 J/L) according to the ventilator setting. An increase of 0.5–0.8 J/L represents a significant additional workload, as WOB in healthy subjects during quiet breathing is about 0.35–0.5 J/L [33, 34]. Furthermore, WOB ≥0.8 J/L has been reported as being associated with weaning failure [35]. Extubating an obese patient after having performed a weaning test without positive pressure could lead to early onset atelectasis if the patient was unable to control for end-expiratory lung volume without PEEP.

**Conclusions**

For the first time the present study reports new insights into respiratory physiology in morbidly obese critically ill candidates to be weaned from the ventilator. These data may be useful for clinicians managing these challenging patients and help make difficult decisions about extubation. We report that either a T piece or a PSV 0 and PEEP 0 cmH₂O test are the trials that predict post-extubation work of breathing in morbidly obese patients. The consequences on mid-term oxygenation and lung aeration, and on the weaning success rate of such weaning tests were, however, not studied.

**Table 4** Inspiratory muscle effort during the five different weaning tests and 20 minutes after extubation

| Test                      | PSV 7 cmH₂O PEEP | PSV 0 cmH₂O PEEP | PSV 7 cmH₂O PEEP | PSV 0 cmH₂O PEEP | T piece | After extubation |
|---------------------------|------------------|------------------|------------------|------------------|---------|-----------------|
| Swing Pes, cmH₂O          | 7.2 ± 5.0*       | 13.4 ± 5.5*      | 12.3 ± 6.3*      | 19.1 ± 7.7       | 19.8 ± 7 | 19.1 ± 5.4      |
| Swing Pdi, cmH₂O          | 8.4 ± 5.5*       | 15.4 ± 5.7*      | 14.2 ± 6.4*      | 21.2 ± 8.1       | 21.7 ± 7 | 20.9 ± 5.5      |
| PTP es, cmH₂O/s/minute    | 141 ± 54*        | 259 ± 84*        | 231 ± 82*        | 346 ± 97         | 332.9 ± 85.9 | 365 ± 87       |
| PTP di, cmH₂O/s/minute    | 157 ± 80*        | 318 ± 113*       | 302 ± 111*       | 451 ± 151        | 439 ± 152 | 465 ± 117       |
| WOB, J/L                  | 0.69 ± 0.31*     | 1.15 ± 0.39*     | 1.09 ± 0.49*     | 1.58 ± 0.57      | 1.57 ± 0.56 | 1.56 ± 0.5      |
| WOB, J/minute             | 7.15 ± 3.5*      | 12.2 ± 6.8*      | 12.4 ± 7.1*      | 17.7 ± 10.2      | 16.8 ± 8.0 | 17.8 ± 9.1      |

Abbreviations: Pdi trans-diaphragmatic pressure, PEEP positive end-expiratory pressure, Pes esophageal pressure, PTPdi trans-diaphragmatic pressure time product, PTPes esophageal pressure time product, PSV pressure support ventilation, WOB work of breathing. *p < 0.001 when compared with after extubation
Additional files

Additional file 1: Supplementary material. (DOCX 44 kb)
Additional file 2: Figure S1, difference in esophageal pressure between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 44 kb)
Additional file 3: Figure S2, difference in the trans-diaphragmatic pressure between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 49 kb)
Additional file 4: Figure S3, difference in the esophageal pressure time product between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 49 kb)
Additional file 5: Figure S4, difference in the trans-diaphragmatic pressure time product between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 47 kb)
Additional file 6: Figure S5, difference in the work of breathing expressed in J/min between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 44 kb)
Additional file 7: Figure S6, difference in the work of breathing expressed in J/min between each test and the post-extubation period. Dashed line represents the absence of difference between the test and the post-extubation period. (JPG 44 kb)

Abbreviations
BMI: Body mass index; CPAP: Continuous positive airway pressure; FI O₂: Inspired oxygen fraction; ICU: Intensive Care Unit; Pdi: Trans-diaphragmatic pressure; PEEP: Positive end-expiration pressure; Pes: Esophageal pressure; Pga: Gastric pressure; PSV: Pressure support ventilation; PTPdi: Trans-diaphragmatic pressure-time product; PTPes: Trans-esophageal pressure-time product; RR: Respiratory rate; SBT: Spontaneous breathing trial; SpO₂: Pulse arterial oxygen saturation; Te: Expiratory time; Ti: Inspiratory time; Ttot: Total cycle duration; VE: Minute ventilation; Vt: Tidal volume; WOB: Work of breathing

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Availability of data and materials
The data that support the findings of this study are available from the corresponding author upon reasonable request.

Authors’ contributions
MM and BJ contributed equally to this work. MM performed the study, analyzed the data and wrote the manuscript. BJ participated in the study design and wrote the manuscript. FG analyzed the data and made critical manuscript revisions. RV analyzed the data and made critical manuscript revisions. YC helped in designing the study, enrolled patients, analyzed the data and made critical manuscript revisions. SM helped in designing the study and extensively corrected the manuscript. AJ performed the statistical analysis and made critical manuscript revisions. MM performed the study, designed and wrote the manuscript. All authors read and approved the final manuscript.

Competing interests
Martin Mahul has nothing to disclose; Boris Jung reports personal fees from Merck (Whitehouse station, NJ) and Astellas (Tokyo, Japan) unrelated to the present study; Fabrice Gallia has nothing to disclose; Nicolas Molinari has nothing to disclose; Audrey De Jong has nothing to disclose; Yannaiël Coisel has nothing to disclose; Rosanna Vaschetto has nothing to disclose; Stefan Matecki has nothing to disclose; Gerald Chaneques has nothing to disclose; Laurent Brochard reports a research contract with Draeger, General Electric and Covidien and honorarium from Draeger unrelated to the present study; Samir Jaber reports personal fees from Maquet, Draeger, Hamilton Medical, Fisher Paykel and Abbott unrelated to the present study.

Consent for publication
Not applicable.

Ethics approval and consent to participate
The present study was approved by the Ethics Committee of the Montpellier Teaching Hospital (2012 A-00294-39, Comité de Protection des Personnes Sud Méditerranée III, Montpellier, France), and registered on clinical trial.gov (reference NCT01616901). All patients provided their written informed consent.

Author details
1Intensive Care Unit, Anaesthesia and Critical Care Department, Saint Elio Teaching Hospital, Centre Hospitalier Universitaire Montpellier, 80 avenue Augustin Fliche, F-34295 Montpellier, Cedex S, France. 2Department of Statistics, University of Montpellier Lapeyronie Hospital, UMR 729 MISTEA, Montpellier, France. 3Anaesthesia and Intensive Care Medicine, Maggiore della Carità Hospital, Novara, Italy. 4Centre National de la Recherche Scientifique (CNRS 9214) – Institut National de la Santé et de la Recherche Médicale (INSERM U-1046), Montpellier University, Montpellier, France. 5Keenan Research Centre, St Michael’s Hospital, Toronto, Canada. 6Interdepartmental Division of Critical Care Medicine, University of Toronto, Toronto, Canada.

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