

“Likely overassistance” during invasive pressure support ventilation in patients in the intensive care unit: a multicentre prospective observational study

Wisam Al-Bassam, Fabian Dade, Michael Bailey, Glenn Eastwood, Eduardo Osawa, Chris Eyeington, James Anesty, George Yi, Jolene Ralph, Nima Kakho, Vishnu Kurup, Elisa Licari, Emma C King, Cameron Knott, Timothy Chimunda, Julie Smith, Ashwin Subramaniam, Mallikarjuna Reddy, Cameron Green, Geoffrey Parkin, Yahya Shehabi and Rinaldo Bellomo

Australia and New Zealand intensive care units (ICUs) admit 150 000 patients per year, of whom 45–58% require mechanical ventilation.^{1,2} Liberation from mechanical ventilation is uncomplicated for most patients;³ however, almost a quarter of patients require extended weaning from the ventilator.⁴ Weaning is a complex, nuanced process that may take up to 50% of the total time a patient spends on ventilatory support.^{4,5}

Pressure support ventilation (PSV) is the most commonly used ventilation mode in Australian ICUs,¹ and it is also commonly used as a weaning mode. It is a spontaneous mode of ventilation that provides titratable inspiratory pressure during a triggered inspiratory effort and a set positive end-expiratory pressure level on expiration.⁶ There is no consensus recommendation for titrating pressure support and there are little data about its use in Australia. However, studies performed in other countries have suggested that variables such as respiratory rate (RR),^{7,8} tidal volume (V_T),⁷⁻⁹ minute volume¹⁰ or rapid shallow breathing index (RSBI)^{11,12} may be used to titrate such pressure support. In particular, there has been concern that inappropriately high levels of pressure support may lead to so called overassistance.

Overassistance during PSV may occur frequently^{13,14} and can lead to excessive V_T and decreased respiratory drive.^{15,16} It may also lead to patient–ventilator dyssynchrony,¹⁵ diaphragmatic atrophy,¹⁷ sleep disorders¹⁸ and ventilator-associated lung injury.¹⁹ Some observational data showed that these side effects may prolong the duration of mechanical ventilation and even mortality.^{20,21} The incidence of such overassistance remains uncertain because it is difficult to diagnose at the bedside. A recent study defined ventilator overassistance as the “occurrence of work of breathing less than 0.3 J/L or 10% or more of ineffective inspiratory effort”.²² Using oesophageal balloon manometry to measure the work of breathing in 27 patients with 211 measurements in a single centre, these investigators found that $RR \leq 17$ breaths/min and/or RSBI (defined as RR/min divided by V_T in litres) ≤ 37 breaths/min/L helped diagnose overassistance at the bedside, with an area

Abstract

Objective: To evaluate the prevalence of “likely overassistance” (categorised by respiratory rate [RR] ≤ 17 breaths/min or rapid shallow breathing index [$RSBI$] ≤ 37 breaths/min/L) during invasive pressure support ventilation (PSV), and the additional prevalence of fixed ventilator settings.

Design: Multicentre prospective observational study of invasive PSV practice in six general Victorian intensive care units with blinding of staff members to data collection.

Patients: At each hospital, investigators collected data between 11 am and 2 pm on all invasive PSV-treated patients on 60 sequential days, excluding weekends and public holidays, between 22 February and 30 August 2017. Each patient was included for maximum of 3 days.

Main results: We studied 231 patients, with a total of 379 observations episodes over the study period. There were 131 patients (56.7%) with at least one episode of $RR \leq 17$ breaths/min; 146 patients (63.2%) with at least one episode of $RSBI \leq 37$ breaths/min/L, and 85 patients (36.8%) with at least one episode of combined $RR \leq 17$ breaths/min and $RSBI \leq 37$ breaths/min/L. Moreover, the total number of observations with “likely overassistance” ($RR \leq 17$ or $RSBI \leq 37$ breaths/min/L) was 178 (47%) and 204 (53.8%), respectively; while for both combined criteria, it was 154 (40.6%). We also found that 10 cmH₂O pressure support was delivered on 210 of the observations (55.4%) and adjusted in less than 25% of observations. Finally, less than half (179 observations) of all PSV-delivered tidal volumes (V_T) were at the recommended value of 6–8 mL/kg predicted body weight (PBW) and more than 20% (79 observations) were at ≥ 10 mL/kg PBW.

Conclusion: In a cohort of Victorian hospitals in Australia, during invasive PSV, “likely overassistance” was common, and the pressure support level was delivered in a standardised and unadjusted manner at 10 cmH₂O, resulting in the frequent delivery of potentially injurious V_T .

under the receiver operating characteristic curve of 0.92 and 0.84, respectively. It also found that $RR \leq 12$ breaths/min had 100% specificity for overassistance, and that overassistance occurred in 37–48% of their study patients.²² However, the prevalence of such overassistance in other centres or health care settings is uncertain. In this regard, there may be considerable heterogeneity in PSV use,²³ and, in particular, there is little information on how PSV is used outside of clinical trial settings, whether it is titrated and whether “likely overassistance” does occur and, if so, how often.

We hypothesised that, in a representative cohort of Australian hospitals, “likely overassistance” (defined as $RR \leq 17$ breaths/min, $RSBI \leq 37$ breaths/min/L, and a combination of both parameters) would occur in at least one-third of such patients or observations, that PSV would be stereotypically set, and that PSV would remain unadjusted in at least half of the treated patients or observations. We conducted a multicentre prospective observational study to test these hypotheses.

Methods

Study design

We conducted a multicentre prospective observational audit of PSV practices in six mixed medical–surgical adult ICUs in Victoria, Australia. Four ICUs were located in tertiary hospitals and two in regional hospitals. This study was classified as low risk, with all information being recorded as part of usual care, and was approved by all ethics committees of the participating institutions with a waiver of informed consent (ethics approval no. LNR/17/Austin/265).

Inclusion criteria

All consecutive adult patients (aged ≥ 18 years) admitted to the participating ICUs during the study period who received invasive mechanical ventilation on PSV mode with either an endotracheal tube, or a nasotracheal tube or a tracheostomy were included in the study.

Table 1. Study patients’ characteristics

Patient characteristics	Values (n = 231)
Male	146 (63.2%)
Age (years), mean (SD)	60 (17.2)
Patient weight, (kg), mean (SD)	83 (22)
Patient height (cm), mean (SD)	169 (11.5)
PBW (kg), mean (SD)	62.7 (15.4)
Elective admission	32 (13.9%)
Medical admission	115 (49.8%)
Premorbid restrictive lung disease	9 (3.9%)
Premorbid obstructive lung disease	41 (17.7%)
Premorbid mixed lung disease	6 (2.6%)
Fluid overload	49 (21.2%)
Acute pulmonary oedema	11 (4.8%)
Pneumonia	68 (29.4%)
Atelectasis	24 (10.4%)
Pleural effusion	65 (28.1%)
APACHE II score, mean (SD)	21.1 (8.34)
Hospital stay before data collection (days), mean (SD)	6.8 (5.9)
ICU stay before data collection (days), mean (SD)	3.5 (5.5)
V_T (mL), mean (SD)	509 (143)
V_T /PBW (mL/kg), mean (SD)	7.8 (6.57)
PEEP, mean (SD)	7.81 (6.57)
Pressure support, median (IQR)	10 (10–10)
Respiratory rate, mean (SD)	17.9 (6.1)
SpO_2 , mean (SD)	96.3 (2.61)
FiO_2 , mean (SD)	30.9 (7.9)
pH, mean (SD)	7.4 (0.073)
Pao_2 , mean (SD)	95.4 (43.5)
$Paco_2$, mean (SD)	40.1 (7.8)
HCO_3^- , mean (SD)	24.9 (4.99)

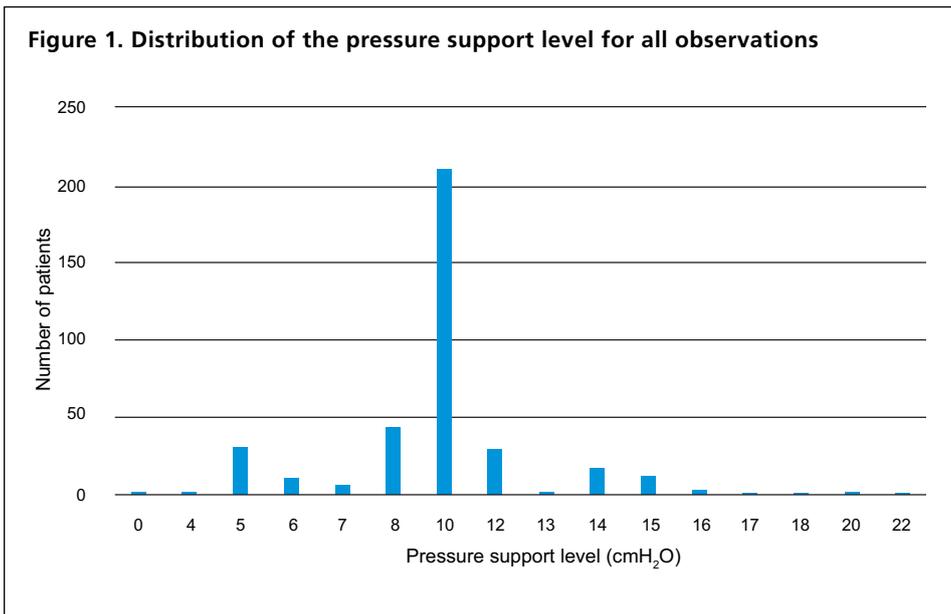
APACHE = Acute Physiology and Chronic Health Evaluation. FiO_2 = fraction of inspired oxygen. HCO_3^- = bicarbonate ion. ICU = intensive care unit. IQR = interquartile range. Pao_2 = arterial partial pressure of oxygen. $Paco_2$ = arterial partial pressure of carbon dioxide. PBW = predicted body weight. PEEP = positive end-expiratory pressure. SD = standard deviation. SpO_2 = oxygen saturation measured by pulse oximetry. V_T = tidal volume.

Exclusion criteria

Patients who received extracorporeal membrane oxygenation, mechanical ventilation but not pressure support ventilation, heart and/or lung transplant, or palliative care were excluded from the study.

Data collection

At each hospital, between 11 am and 2 pm, investigators collected data on patients meeting the inclusion criteria

Figure 1. Distribution of the pressure support level for all observations

Data analysis

All data were initially assessed for normality. Group comparisons were performed using χ^2 test for equal proportion (or Fisher exact test when numbers were small), Student *t* test for normally distributed data, and Wilcoxon rank sum otherwise, with results reported as numbers (%), means \pm standard deviation or median (interquartile range [IQR]), respectively. To increase the robustness of this analysis, a two-sided $P = 0.01$ has been chosen to indicate statistical significance. All analyses were performed using SAS version 9.4 (SAS Institute, Cary, NC, USA).

on 60 sequential days (excluding weekends and public holidays) between 22 February and 30 August 2017. Each patient was included for maximum of 3 days. The variables recorded were all available at the bedside or on the computer-based ICU system. We only used data pertaining to the patient's index (first) ICU admission. Using existing ICU-based electronic databases and an electronic medical record review, we recorded all arterial blood gas results and documented chest x-ray reports for all patients involved in the study.

The attending clinical staff, including all medical and nursing staff, were blinded to the conduct and purpose of the study to avoid changes in practice. Data collection included patients' demographics, APACHE (Acute Physiology and Chronic Health Evaluation) II score, past medical history and admitting diagnosis, ventilator settings and variables.

Outcomes

The primary outcome was the prevalence of "likely overassistance" (defined as presence of $RR \leq 17$ breaths/min and $RSBI \leq 37$ breaths/min/L), as previously described.²⁰

The secondary outcomes were the percentage of patients and/or observations with the level of pressure support set at a fixed unadjusted value, defined by the absence of any changes during the observation period. The other secondary outcome was the percentage of patients and observations with $V_T > 8$ mL/kg predicted body weight (PBW) and > 10 mL/kg PBW. PBW was calculated using the criteria from the National Institutes of Health–National Heart, Lung, and Blood Institute Acute Respiratory Distress Syndrome (ARDS) Network.²⁴

Results

Patient characteristics

We studied 231 patients, including a total of 379 observation points in six ICUs over the study period. Overall, 146 patients were male (63%), the mean patient age was 60 ± 17 years and the average PBW was 62.7 ± 15.4 kg. The average APACHE II score was 21.1 ± 8.3 , and patients were assessed at a median of Day 3 (IQR, 1–7) of ICU stay (Table 1). Nine patients had pre-morbid restrictive lung disease, and 41 had chronic obstructive pulmonary disease. One in five patients was diagnosed as having fluid overload, and one in four as having pneumonia (Table 1). The median level of pressure support was 10 cmH₂O (IQR, 10–10) (Table 1). The total number of all observations in the study patients was 379, and a pressure support level 10 cmH₂O was applied in 210 patients (55%) (Figure 1).

Patients with "likely overassistance"

Overall, of 231 patients, 131 (56.7%) had the first marker of "likely overassistance" (at least one episode of $RR \leq 17$ breaths/min) (online Appendix, Table S1, available at cicm.org.au/Resources/Publications/Journal). These patients had significantly higher V_T and higher V_T /PBW ratios, and a significantly lower RR. One-hundred and forty-six patients (63.2%) had the second marker of "likely overassistance" (at least one episode of $RSBI \leq 37$ breaths/min/L). They were more likely to be male, had higher actual and predicted body weight and higher V_T and V_T /PBW ratios (online Appendix, Table S2).

There were 85 patients (36.8%) with both markers of "likely overassistance" (online Appendix, Table S3). Their

Table 2. Summary of the total observations for different respiratory variables

Variable	Total number of observations	Number (%)
RR \leq 17 (breaths/min)	379	178 (47%)
RR \leq 12 (breaths/min)	379	71 (18.7%)
RSBI \leq 37 (breaths/min/L)	379	204 (53.8%)
RR \leq 17 (breaths/min) and RSBI \leq 37 (breaths/min/L)	379	154 (40.6%)
V_T /PBW $>$ 8 (mL/kg)	378	172 (45.5%)
V_T /PBW $>$ 10 (mL/kg)	378	79 (20.9%)

PBW = predicted body weight. RR = respiratory rate. RSBI = respiratory shallow breathing index. V_T = tidal volume.

support level was decreased only in 13 observations (18.3%) and, interestingly, 11 observations (15.5%) had their pressure support level increased despite V_T /PBW $>$ 8 mL/kg (Table 3).

Thirty-five out of 148 observations (23.6%) remained ventilated with $V_T >$ 10mL/kg PBW the subsequent day. Of these, 17 observations (48.6%) had no change in pressure support level, and 11 (31.4%) remained at a pressure support level 10 cmH₂O (Table 3).

Discussion

Key findings

In a multicentre prospective observational study of current PSV practice in six Australian hospitals, we found that “likely overassistance” (defined as RR \leq 17 breaths/min or RSBI \leq 37 breaths/min/L) was widespread and involved more than 50% of all patients and observations. Moreover, we found that a standard pressure support level of 10 cmH₂O was delivered during more than half of the observation episodes. In addition, we also found that less than half of the total observations were at the recommended V_T /PBW ratio of 6–8 mL/kg, and that more than 20% of observations were at a V_T /PBW \geq 10 mL/kg (Table 2). Finally, the subsequent day, most patients had no adjustment in pressure support levels despite “likely overassistance” and high V_T delivery, even for patients with RR \leq 12 breaths/min or V_T /PBW \geq 10 mL/kg (Table 3).

Relationship to previous studies

PSV is one of the most commonly used ventilation strategies in Australia, but the optimum pressure support level is not well defined or studied. This is problematic because an unadjusted pressure support level may result in high V_T and overassistance, both of which can cause harm, such as ventilator-induced lung injury,^{25,26} prolonged ventilation,¹⁶ hyperinflations,²⁶ diaphragmatic atrophy¹⁷ and patient-ventilator dyssynchrony.^{15,27}

Despite the wide use of PSV,¹ overassistance has not been studied widely — the majority of the studies have evaluated underassistance.^{13,14} Pletsch-Assuncao and colleagues²² found a rate of overassistance between 37% and 48% at baseline in a cohort of 27 patients with 211 observations. Our findings are similar, identifying “likely overassistance” in 40–53% of all observations. This is despite several observations in the literature that suggest harm from overassistance. For example, Leung et al²⁸ compared associated ventilator modes on triggering, patient effort and

characteristics replicated those in each of the two separate “likely overassistance” groups, including a shorter time in the ICU and hospital before the first measurement.

Observations with “likely overassistance”

The study had a total of 379 observations episodes (Table 2). The total number of observations with a marker of “likely overassistance” (RR \leq 17 or RSBI \leq 37 breaths/min/L) was 178 (46.9%) and 204 (53.8%), respectively; while for both combined criteria it was 154 (40.6%) (Table 2). Moreover, 71 observations (18.7%) had RR \leq 12 breaths/min.

Response to “likely overassistance”

There were 148 observations in which a follow-up observation was available the next day (Table 3). Sixty-one observations had RR \leq 17 breaths/min, 36 (59%) had no changes in their pressure support levels and 28 (45.9%) remained at an unadjusted pressure support level of 10 cmH₂O the next day. Moreover, 11 (18%) had their pressure support level increased, and only 14 (22.9%) had a decrease in pressure support levels. Among those with RSBI \leq 37 breaths/min/L at the first observation and those with both “likely overassistance” markers, the findings were similar (Table 3) (online Appendix, Figure S1). Finally, such pattern of limited adjustment applied even to patients with RR \leq 12 breaths/min (Table 3).

Tidal volume to predicted body weight ratio

Almost half of the observation episodes (172/378 observations) had V_T in excess of 8 mL/kg PBW (Table 2). A total of 148 observations were followed up the next day; 71 of these observations (48%) had $V_T >$ 8mL/kg PBW the next day. Forty-seven of 71 observations (66.2%) continued to have the same level of pressure support the subsequent day, with V_T levels remaining above 8 mL/kg PBW. Pressure

Table 3. Summary of the data for the next day follow-up

Respiratory variables	Total observations (n = 148)	Percentage of observations			
		Pressure support level decreased	Pressure support level unchanged	Pressure support level increased	Pressure support level remained at 10 cmH ₂ O
RR ≤ 17 (breaths/min)	61 (41.2%)	14 (22.9%)	36 (59%)	11 (18%)	28 (45.9%)
RR ≤ 12 (breaths/min)	22 (14.9%)	4 (18.1%)	14 (63.6%)	4 (18.1%)	10 (45.4%)
RSBI ≤ 37 (breaths/min/L)	70 (47.3%)	13 (18.6%)	45 (64.3%)	12 (17.1%)	32 (45.7%)
RR ≤ 17 (breaths/min) and RSBI ≤ 37 (breaths/min/L)	52 (35.1%)	12 (23%)	32 (61.5%)	8 (15.3%)	25 (48%)
V _T /PBW > 8 (mL/kg)	71 (48%)	13 (18.3%)	47 (66.2%)	11 (15.5%)	31 (43.7%)
V _T /PBW > 10 (mL/kg)	35 (23.6%)	9 (25.7%)	17 (48.6%)	9 (25.7%)	11 (31.4%)

PBW = predicted body weight. RR = respiratory rate. RSBI = respiratory shallow breathing index. V_T = tidal volume.

dyspnoea and showed that adding high pressure support increased the rate of ineffective triggering and wasted inspiratory effort. Moreover, high levels of pressure support produced large V_T and prolonged inspiratory time.²⁹ Large V_T can decrease the neural inspiratory time and inspiratory effort.³⁰ Thille and colleagues¹⁵ showed that aiming for 6 mL/kg during PSV can eliminate ineffective triggering and reduce patient–ventilator dyssynchrony. A meta-analysis performed by Serpa Neto et al³¹ showed that among patients without ARDS lung-protective ventilation with lower V_T could have better outcomes.³¹ PBW is more reliable than actual body weight to estimate lung size because actual body weight can produce excessive V_T in patients with obesity and inadequate V_T in underweight patients. In mandatory mode of ventilations, patients with ARDS who are ventilated > 8 mL/kg PBW³² can have ventilator-induced lung injury. It is possible that PSV at similarly high volumes may also be injurious.⁹ A retrospective analysis study by Amato et al,³³ indicated that patients with ARDS who are on mandatory mode of ventilation benefit from reductions in V_T only if associated with a decrease in driving pressure (calculated by plateau pressure minus driving pressure), as low driving pressure can reduce the stress on lungs with low compliance. Similarly, in PSV, a decrease in driving pressure by lowering pressure support level in order to achieve a low V_T might also lead to a decrease in lung stress in spontaneously ventilated patients.²²

Implications of study findings

Our findings imply that in patients treated in Victorian ICUs “likely overassistance” is common. PSV is delivered in a stereotypical manner at 10 cmH₂O. Finally, they imply that despite “likely overassistance” and high V_T/PBW ratios in the likely injurious range, the level of PSV is typically unadjusted toward a safer and more physiological level.

Strengths and limitations

To our knowledge, this is the only multicentre study to address pressure support levels and the occurrence of “likely overassistance”, as recently defined.²⁰ Our study included six different ICUs, making our findings likely representative of both Victoria and Australia. We studied PSV strategies in ICUs ranging from tertiary to regional hospitals which included trauma, liver transplant, cardiothoracic, neurosurgical and medical ICU patients. The total number of patients included in the study was > 200 and the number of observations almost 400, thus likely to reliably represent overall daily PSV practice. Finally, all ICU staff were blinded to the purpose and conduct of the study at the time of the data collection, preventing a Hawthorne effect.

Our study has some limitations. It was performed in Victorian hospitals and may not apply to hospitals in other countries. However, our results are very similar to the study by Pletsch-Assuncao and colleagues²² in Brazil, suggesting that our findings may apply beyond the Australian context. Moreover, the study hospitals had a twice daily ward round by an ICU specialist, giving ample opportunity for adjustment of pressure support levels. The data collection was not based on continuous data recordings and, for pragmatic reasons, was performed as a single data collection point per day. Nevertheless, the follow-up data showed that the changes in pressure support level occurred uncommonly and the rate of “likely overassistance” remained overall unchanged. For “likely overassistance” identification, we relied on the findings of a previous study.²⁰ Therefore, we used the qualifier “likely” because we did not conduct detailed invasive assessment of work of breathing. However, our findings are similar to those obtained with such measurements, have face validity and, even when our criteria were made stricter by using RR ≤ 12 breaths/min, and more specific by using both RR and RSBI, the results

were not significantly different. The group of patients with RR > 17 breaths/min, RSBI > 37 breaths/min/L and the combination of both markers (online Appendix, Table S2, Table S3 and Figure S1) had an average stay in the ICU before data collection of several days more than the patients with “likely overassistance”. However, APACHE score, premorbid conditions and diagnosis were similar. Hence, the longer stay of patients before data collection is unlikely to have a significant implication on the study results.

Conclusion

In a multicentre prospective observational study in Australia, we found that, during PSV, “likely overassistance” was widespread, a standard level of pressure support of 10 cmH₂O was delivered most of the time and, with little follow-up adjustment, despite low RR and low RSBI, most observations were at V_T/PBW ratios in the potentially injurious range, also with little adjustment. These observations raise concerns about the quality and safety of invasive PSV in Australian ICUs.

Competing interests

None declared.

Author details

Wisam Al-Bassam¹
 Fabian Dade²
 Michael Bailey³
 Glenn Eastwood⁴
 Eduardo Osawa⁴
 Chris Eyeington⁴
 James Anesty²
 George Yi²
 Jolene Ralph²
 Nima Kakho⁵
 Vishnu Kurup⁵
 Elisa Licari⁶
 Emma C King^{6,7}
 Cameron Knott^{4,8}
 Timothy Chimunda⁸
 Julie Smith⁸
 Ashwin Subramaniam^{7,9}
 Mallikarjuna Reddy^{9,10}
 Cameron Green⁹
 Geoffrey Parkin⁷
 Yahya Shehabi¹
 Rinaldo Bellomo^{3,4}

- 1 Monash Medical Centre, Melbourne, VIC, Australia.
- 2 Royal Melbourne Hospital, Melbourne, VIC, Australia.
- 3 Australian and New Zealand Intensive Care Research Centre, Monash University, Melbourne, VIC, Australia.
- 4 Austin Health, Melbourne, VIC, Australia.
- 5 Barwon Health, Geelong, VIC, Australia.
- 6 Alfred Health, Melbourne, VIC, Australia.
- 7 Monash University, Melbourne, VIC, Australia.
- 8 Bendigo Health, Bendigo, VIC, Australia.
- 9 Frankston Hospital, Frankston, VIC, Australia.
- 10 Peninsula Health, Frankston, VIC, Australia.

Correspondence: Wisam.AlBassam@monashhealth.org.au

Participating institutions

- Department of Intensive Care, Austin Health, Melbourne, VIC, Australia.
- Department of Intensive Care, Royal Melbourne Hospital, Melbourne, VIC, Australia.
- Department of Intensive Care, Alfred Hospital, Melbourne, VIC, Australia.
- Department of Intensive Care, Bendigo Health, Bendigo, VIC, Australia.
- Department of Intensive Care, Frankston Hospital, Frankston, VIC, Australia.
- Department of Intensive Care, Barwon Health, Geelong, VIC, Australia.
- Department of Intensive Care, Monash Medical Centre, Melbourne, VIC, Australia.
- Australian and New Zealand Intensive Care Research Centre, Monash University, Melbourne, VIC, Australia.

References

- 1 Rose L, Presneill JJ, Johnston L, et al. Ventilation and weaning practices in Australia and New Zealand. *Anaesth Intensive Care* 2009; 37: 99-107.
- 2 Berney SC, Harrold M, Webb SA, et al. Intensive care unit mobility practices in Australia and New Zealand: a point prevalence study. *Crit Care Resusc* 2013; 15: 260-5.
- 3 Esteban A, Ferguson ND, Meade MO, et al. Evolution of mechanical ventilation in response to clinical research. *Am J Respir Crit Care Med* 2008; 177: 170-7.
- 4 Boles JM, Bion J, Connors A, et al. Weaning from mechanical ventilation. *Eur Respir J* 2007; 29: 1033-56.
- 5 Esteban A, Alía I, Ibañez J, et al. Modes of mechanical ventilation and weaning. A national survey of Spanish hospitals. The Spanish Lung Failure Collaborative Group. *Chest* 1994; 106: 1188-93.

- 6 Pardo M. Mechanical and noninvasive ventilation. In: Parsons PE, Weiner-Kronish JP, editors. *Critical care secrets*. 5th ed. St Louis, MO: Elsevier Mosby, 2013; pp 58-62.
- 7 Alberti A, Gallo F, Fongaro A, et al. P0.1 is a useful parameter in setting the level of pressure support ventilation. *Intensive Care Med* 1995; 21: 547-53.
- 8 Banner MJ, Kirby RR, Kirton OC, et al. Breathing frequency and pattern are poor predictors of work of breathing in patients receiving pressure support ventilation. *Chest* 1995; 108: 1338-44.
- 9 Writing Group for the PREVENT Investigators; Simonis FD, Serpa Neto A, Binnekade JM, et al. Effect of a low vs intermediate tidal volume strategy on ventilator-free days in intensive care unit patients without ARDS: A randomized clinical trial. *JAMA* 2018; 320: 1872-80.
- 10 Van de Graaff WB, Gordey K, Dornseif SE, et al. Pressure support. Changes in ventilatory pattern and components of the work of breathing. *Chest* 1991; 100: 1082-9.
- 11 Samanta S, Singh RK, Baronia AK, et al. Diaphragm thickening fraction to predict weaning — a prospective exploratory study. *J Intensive Care* 2017; 5: 62.
- 12 Yang KL, Tobin MJ. A prospective study of indexes predicting the outcome of trials of weaning from mechanical ventilation. *N Engl J Med* 1991; 324: 1445-50.
- 13 Ducharme-Crevier L, Du Pont-Thibodeau G, Emeriaud G. Interest of monitoring diaphragmatic electrical activity in the pediatric intensive care unit. *Crit Care Res Pract* 2013; 2013: 384210.
- 14 Emeriaud G, Larouche A, Ducharme-Crevier L, et al. Evolution of inspiratory diaphragm activity in children over the course of the PICU stay. *Intensive Care Med* 2014; 40: 1718-26.
- 15 Thille AW, Cabello B, Galia F, et al. Reduction of patient-ventilator asynchrony by reducing tidal volume during pressure-support ventilation. *Intensive Care Med* 2008; 34: 1477-86.
- 16 de Wit M, Miller KB, Green DA, et al. Ineffective triggering predicts increased duration of mechanical ventilation. *Crit Care Med* 2009; 37: 2740-5.
- 17 Hudson MB, Smuder AJ, Nelson WB, et al. Both high level pressure support ventilation and controlled mechanical ventilation induce diaphragm dysfunction and atrophy. *Crit Care Med* 2012; 40: 1254-60.
- 18 Parthasarathy S, Tobin MJ. Effect of ventilator mode on sleep quality in critically ill patients. *Am J Respir Crit Care Med* 2002; 166: 1423-9.
- 19 Acute Respiratory Distress Syndrome Network, Brower RG, Matthay MA, Morris A, et al. Ventilation with lower tidal volumes as compared with traditional tidal volumes for acute lung injury and the acute respiratory distress syndrome. *N Engl J Med* 2000; 342: 1301-8.
- 20 Blanch L, Villagra A, Sales B, et al. Asynchronies during mechanical ventilation are associated with mortality. *Intensive Care Med* 2015; 41: 633-41.
- 21 Stewart NI, Jagelman TA, Webster NR. Emerging modes of ventilation in the intensive care unit. *Br J Anaesth* 2011; 107: 74-82.
- 22 Pletsch-Assuncao R, Caleffi Pereira M, Ferreira JG, et al. Accuracy of invasive and noninvasive parameters for diagnosing ventilatory overassistance during pressure support ventilation. *Crit Care Med* 2018; 46: 411-7.
- 23 Australian and New Zealand Intensive Care Society Centre for Outcome and Resource Evaluation. Adult Patient Database activity report 2015–2016. Melbourne: ANZICS; 2016. <https://www.anzics.com.au/wp-content/uploads/2018/08/ANZICS-CORE-APD-Activity-Report-2015-16.pdf> (viewed Dec 2018).
- 24 Linares-Perdomo O, East TD, Brower R, et al. Standardizing predicted body weight equations for mechanical ventilation tidal volume settings. *Chest* 2015; 148: 73-78.
- 25 Brower RG, Matthay MA, Morris A, et al. Ventilation with lower tidal volumes as compared with traditional tidal volumes for acute lung injury and the acute respiratory distress syndrome. *N Engl J Med* 2000; 342: 1301-8.
- 26 Dreyfuss D, Soler P, Basset G, Saumon G. High inflation pressure pulmonary edema. Respective effects of high airway pressure, high tidal volume, and positive end-expiratory pressure. *Am Rev Respir Dis* 1988; 137: 1159-64.
- 27 Thille AW, Rodriguez P, Cabello B, et al. Patient-ventilator asynchrony during assisted mechanical ventilation. *Intensive Care Med* 2006; 32: 1515-22.
- 28 Leung P, Jubran A, Tobin MJ. Comparison of assisted ventilator modes on triggering, patient effort, and dyspnea. *Am J Respir Crit Care Med* 1997; 155: 1940-8.
- 29 Jubran A, Van de Graaff WB, Tobin MJ. Variability of patient-ventilator interaction with pressure support ventilation in patients with chronic obstructive pulmonary disease. *Am J Respir Crit Care Med* 1995; 152: 129-36.
- 30 Clark FJ, von Euler C. On the regulation of depth and rate of breathing. *J Physiol* 1972; 222: 267-95.
- 31 Serpa Neto A, Cardoso SO, Manetta JA, et al. Association between use of lung-protective ventilation with lower tidal volumes and clinical outcomes among patients without acute respiratory distress syndrome: a meta-analysis. *JAMA* 2012; 308: 1651-9.
- 32 Antonelli M, Azoulay E, Bonten M, et al. Year in review in intensive care medicine 2010: III. ARDS and ALI, mechanical ventilation, noninvasive ventilation, weaning, endotracheal intubation, lung ultrasound and paediatrics. *Intensive Care Med* 2011; 37: 394-410.
- 33 Amato MB, Meade MO, Slutsky AS, et al. Driving pressure and survival in the acute respiratory distress syndrome. *N Engl J Med* 2015; 372: 747-55.

Appendix

This appendix was part of the submitted manuscript and has been peer reviewed. It is posted as supplied by the authors.

Table S1. Patients characteristics for RR less or equal to 17 versus more than 17

	RR ≤ 17 ¹ (n = 131)	RR > 17 ² (n = 100)	p value
Patient Characteristics	(% or SD) ³	(% or SD) ³	
Male	87 (66.4%)	59 (59%)	0.25
Age	60 (17)	60 (17.5)	0.99
Patient weight (KG)	83.8 (23.1)	82 (20.4)	0.55
Patient height (cm)	170 (10.7)	168 (12.4)	0.25
PBW⁴ (KG)	64.3 (11.1)	60.8 (19.5)	0.09
Elective admission	81 (13.7%)	14 (14%)	0.96
Medical admission	62 (47.3%)	53 (53%)	0.39
Premorbid restrictive lung disease	5 (3.8%)	4 (4%)	0.95
Premorbid obstructive lung disease	21 (16%)	20 (20%)	0.43
Premorbid mixed lung disease	4 (3.1%)	2 (2%)	0.7
Fluid overload	26 (20%)	23 (23%)	0.58
APO⁵	7 (5.4%)	4 (4%)	0.63
Pneumonia	30 (23.1%)	38 (38%)	0.014
Atelectasis	15 (11.5%)	9 (9%)	0.53
Pleural effusion by CXR	40 (30.8%)	25 (25%)	0.36
APACHE II score	20.1 (8.06)	22.4 (8.55)	0.035
Hospital days before data collection	6.6 (13.3)	7.7 (8.35)	0.01
ICU days before data collection	2.7 (4.2)	5.2 (7.4)	0.001
VT⁶ (ml)	546 (147)	461 (123)	<0.0001
VT/PBW⁷ (ml/kg)	8.7 (2.5)	7.8 (5.7)	<0.0001
PEEP⁸	8 (8.46)	7.6 (2.47)	0.65
PS⁹	10 [8-10]	10 [10-10]	0.08
RR¹⁰	14.3 (4.34)	22.7 (4.61)	<0.0001
SpO₂¹¹	96.5 (2.43)	96 (2.82)	0.14
FiO₂¹²	30.4 (7.7)	31.5 (8.2)	0.3
pH¹³	7.39 (0.076)	7.42 (0.066)	0.003
PaO₂¹⁴	97.3 (47.9)	92.9 (37.2)	0.46
PaCO₂¹⁵	41.2 (7.5)	38.8 (8.2)	0.015
HCO₃¹⁶	24.7 (4.8)	25.1 (5.25)	0.53

- 1- (% or SD) = Percentage or standard deviation
- 2- RR ≤ 17 = Respiratory rate less or equal to 17
- 3- RR > 17 = Respiratory rate more than 17
- 4- PBW = Predicted body weight
- 5- APO = Acute pulmonary oedema
- 6- VT = Tidal volume
- 7- VT/PBW = Tidal volume on predicted body weight
- 8- PEEP = Positive end expiratory pressure
- 9- PS = Pressure support
- 10- RR = Respiratory rate
- 11- SpO₂ = Oxygen saturation
- 12- FIO₂ = Fractional of inspired oxygen
- 13- pH = Scale of acidity
- 14- PaO₂ = Partial pressure of oxygen
- 15- PaCO₂ = Partial pressure of carbon dioxide
- 16- HCO₃ = Bicarbonate ion

Table S2. Patient characteristics for RSBI less than or equal to 37 versus RSBI more than 37

Patient characteristics	RSBI \leq 37 ¹ (n=146) (% or SD) ³	RSBI > 37 ² (n=85) (% or SD) ³	pvalue
Male	108 (74%)	38 (44.7%)	<0.0001
Age	59.1 (16.5)	61.7 (18.3)	0.26
Patient weight (KG)	86 (22.6)	77.8 (19.9)	0.007
Patient Height (cm)	171 (10.4)	166 (12.5)	<0.0001
PBW ⁴ (KG)	65.7 (10.6)	57.6 (20.2)	<0.0001
Elective admission	23 (15.8%)	9 (10.6%)	0.27
Medical admission	72 (49.3%)	43 (50.6%)	0.85
Premorbid restrictive lung disease	5 (3.4%)	4 (4.7%)	0.64
Premorbid obstructive lung disease	23 (15.8%)	18 (21.2%)	0.30
Premorbid mixed lung disease	4 (2.7%)	2 (2.4%)	1.00
Fluid overload	29 (19.9%)	20 (23.8%)	0.48
APO ⁵	6 (4.1%)	5 (6%)	0.53
Pneumonia	35 (24%)	33 (39.3%)	0.01
Atelectasis	13 (8.9%)	11 (13.1%)	0.32
Pleural effusion	40 (27.6%)	25 (29.8%)	0.73
APACHE II score	20.5 (8.23)	22.2 (8.47)	0.13
Hospital days before data collection	5.9 (10.4)	8.3 (13)	0.005
ICU days before data collection	3.1 (5.6)	5 (6.3)	<0.0001
VT ⁶ (ml)	575 (129)	397 (84.2)	<0.0001
VT/PBW ⁷	9 (2.5)	7.1 (6)	<0.0001
PEEP ⁸	7.92 (8.09)	7.64 (2.28)	0.75
PS ⁹	10 [8-10]	10 [10-12]	0.001
RR ¹⁰	15 (4.52)	22.9 (5.15)	<0.0001
SpO ₂ ¹¹	96.4 (2.43)	96.2 (2.91)	0.52
FiO ₂ ¹²	3.05 (7.9)	31.4 (8.1)	0.41
pH ¹³	7.4 (0.074)	7.41 (0.071)	0.13
PaO ₂ ¹⁴	94.7 (45.5)	96.5 (40)	0.76
PaCO ₂ ¹⁵	40 (7.77)	40.3 (8.19)	0.80
HCO ₃ ¹⁶	24.4 (4.79)	25.6 (5.27)	0.08

1- (% or SD) = Percentage or standard deviation

2- RSBI \leq 37 = Rapid shallow breathing index less or equal to 37

3- RSBI > 17 = Rapid shallow breathing index more than 17

4- PBW = Predicted body weight

5- APO = Acute pulmonary oedema

6- VT = Tidal volume

7- VT/PBW = Tidal volume on predicted body weight

8- PEEP = Positive end expiratory pressure

9- PS = Pressure support

10- RR = Respiratory rate

11- SpO₂ = Oxygen saturation

12- FiO₂ = Fractional of inspired oxygen

13- pH = Scale of acidity

14- PaO₂ = Partial pressure of oxygen

15- PaCO₂ = Partial pressure of carbon dioxide

16- HCO₃ = Bicarbonate ion

Table S3. Patients characteristics for combination of RR less or equal to 17 and RSBI less or equal to 37 versus RR more than 17 and RSBI more than 37

Patients characteristics	RR ≤ 17and RSBI ≤37¹ (% or SD) ³	RR > 17and RSBI >37² (% or SD)	pvalue
Male	85 (73.3%)	61 (53%)	0.001
Age	59.8 (16.4)	60.2 (18)	0.87
Patient weight (KG)	85.8 (23.4)	80.3 (20.2)	0.06
Patient height (cm)	171 (10.5)	167 (12.2)	0.01
PBW⁴ (KG)	65.3 (10.8)	60.2 (18.6)	0.005
Elective admission	17 (14.7%)	15 (13%)	0.72
Medical admission	55 (47.4%)	60 (52.2%)	0.47
Premorbid restrictive lung disease	5 (4.3%)	4 (3.5%)	0.73
Premorbid obstructive lung disease	18 (15.5%)	23 (20%)	0.37
Premorbid mixed lung disease	4 (3.4%)	2 (1.7%)	0.68
Fluid overload	23 (19.8%)	26 (22.8%)	0.58
APO⁵	6 (5.2%)	5 (4.4%)	0.78
Pneumonia	26 (22.4%)	42 (36.8%)	0.02
Atelectasis	11 (9.5%)	13 (11.4%)	0.63
Pleural effusion	33 (28.7%)	32 (28.1%)	0.92
Apache II score	20.2 (8.29)	22 (8.33)	0.10
Hospital days before data collection	5.8 (10.6)	7.8 (12.2)	0.008
ICU days before data collection	2.7 (4.4)	4.8 (6.9)	0.001
VT⁶ (ml)	571 (136)	447 (121)	<0.0001
VT/PBW⁷ (ml/kg)	9 (2.5)	7.6 (5.3)	<0.0001
PEEP⁸	8 (8.95)	7.54 (2.46)	0.53
PS⁹	10 [8-10]	10 [10-10]	0.09
RR¹⁰	13.8 (4.33)	22 (4.77)	<0.0001
SpO₂¹¹	96.6 (2.48)	96.1 (2.73)	0.14
FiO₂¹²	30.8 (7.9)	30.9 (8)	0.94
pH¹³	7.39 (0.076)	7.42 (0.068)	0.004
PaO₂¹⁴	95.3 (46.2)	95.5 (40.8)	0.98
PaCO₂¹⁵	41.1 (7.7)	39.2 (7.9)	0.06
HCO₃¹⁶	24.6 (4.91)	25.1 (5.08)	0.42

1- (% or SD) = Percentage or standard deviation

2- RR ≤ 17 and RSBI ≤ 37 = RR less or equal to 17 and rapid shallow breathing index less or equal to 37

3- RR > 17 and RSBI > 37= RR more than 17 and rapid shallow breathing index more than 17

4- PBW = Predicted body weight

5- APO = Acute pulmonary oedema

6- VT = Tidal volume

7- VT/PBW = Tidal volume on predicted body weight

8- PEEP = Positive end expiratory pressure

9- PS = Pressure support

10- RR = Respiratory rate

11- SpO₂ = Oxygen saturation

12- FIO₂ = Fractional of inspired oxygen

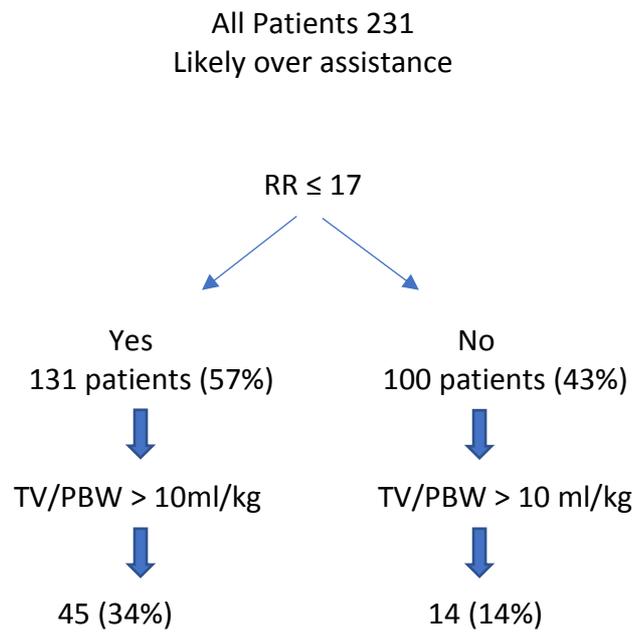
13- pH = Scale of acidity

14- PaO₂ = Partial pressure of oxygen

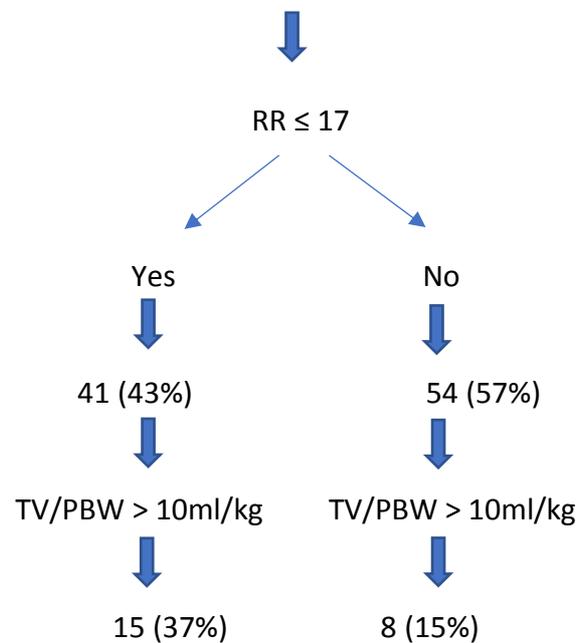
15- PaCO₂= Partial pressure of carbon dioxide

16- HCO₃ = Bicarbonate ion

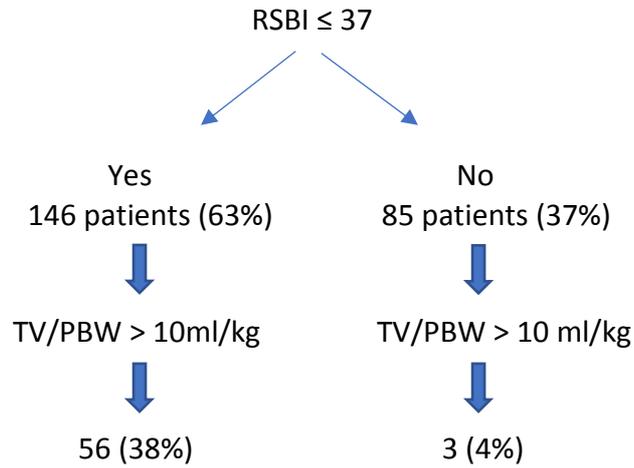
Appendix Figure S1



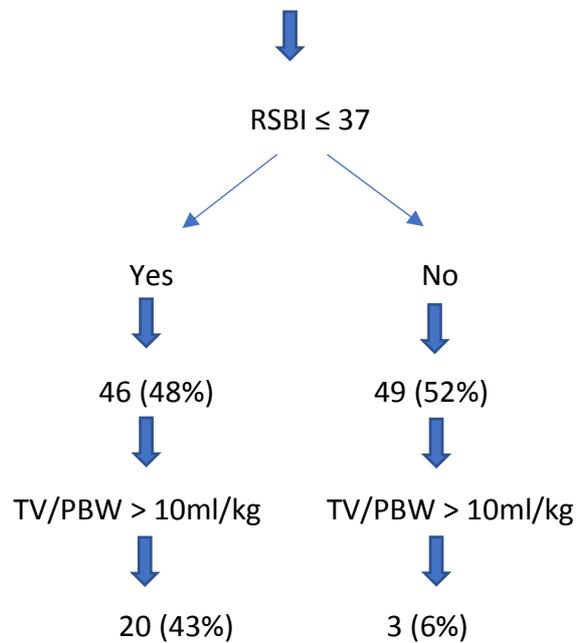
Following day data collection
Number of patients followed up on day 2: n=95 (41%)
Likely over assistance



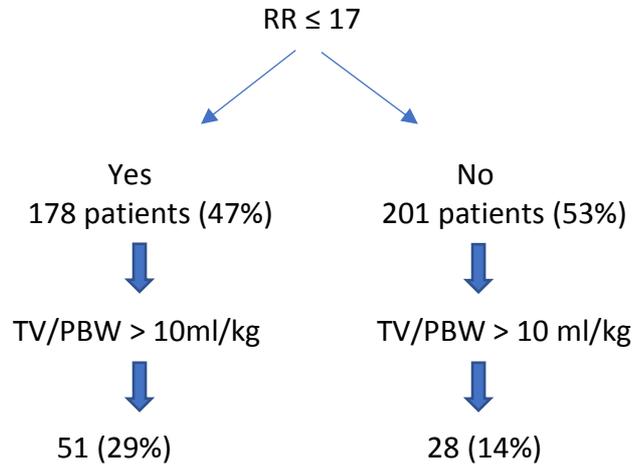
All Patients 231
Likely over assistance



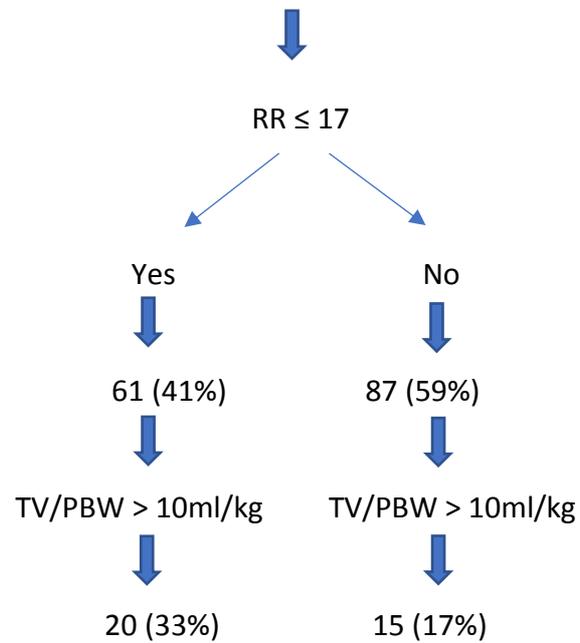
Following day data collection
Number of patients followed up on day 2: 95 (41%)
Likely over assistance



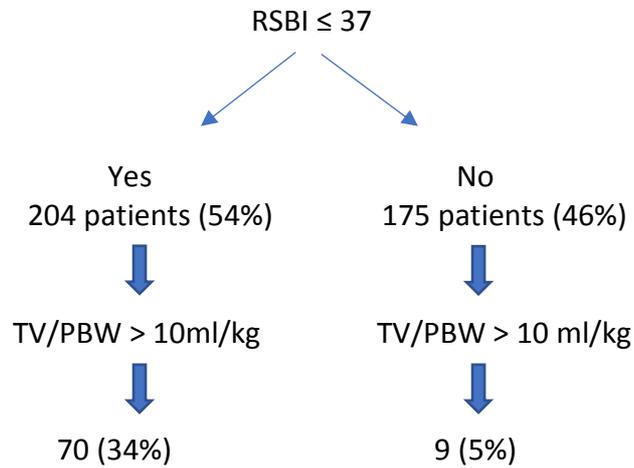
All observations 379
Likely over assistance



Following day data collection
Number of observations followed up next day (148)
Likely over assistance



All observations 379
Likely over assistance



Following day data collection
Number of observations followed up next day (148)
Likely over assistance

