

HHS Public Access

Author manuscript *Anaesthesia.* Author manuscript; available in PMC 2020 August 13.

Published in final edited form as: *Anaesthesia*. 2019 September ; 74(9): 1165–1174. doi:10.1111/anae.14742.

The development and validation of the Score for the Prediction of Postoperative Respiratory Complications (SPORC-2) to predict the requirement for early postoperative tracheal reintubation: a hospital registry study

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Abstract

Postoperative pulmonary complications are associated with an increase in mortality, morbidity and healthcare utilisation. The Agency for Healthcare Research and Quality recommends risk assessment for postoperative respiratory complications in patients undergoing surgery. In this hospital registry study of adult patients undergoing non-cardiac surgery between 2005 and 2017 at two independent healthcare networks, a prediction instrument for early postoperative tracheal reintubation was developed and externally validated. This was based on the development of the Score for Prediction Of Postoperative Respiratory Complications. For predictor selection, stepwise

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backward logistic regression and bootstrap resampling were applied. Development and validation cohorts were represented by 90,893 patients at Partners Healthcare and 67,046 patients at Beth Israel Deaconess Medical Center, of whom 699 (0.8%) and 587 (0.9%) patients, respectively, had their tracheas reintubated. In addition to five pre-operative predictors identified in the Score for Prediction Of Postoperative Respiratory Complications, the final model included seven additional intra-operative predictors: early post-tracheal intubation desaturation; prolonged duration of surgery; high fraction of inspired oxygen; high vasopressor dose; blood transfusion; the absence of volatile anaesthetic use; and the absence of lung-protective ventilation. The area under the receiver operating characteristic curve for the new score was significantly greater than that of the original Score for Prediction Of Postoperative Respiratory Complications (0.84 [95%CI 0.82–0.85] vs. 0.76 [95%CI 0.75–0.78], respectively; p < 0.001). This may allow clinicians to develop and implement strategies to decrease the risk of early postoperative tracheal re-intubation.

Keywords

clinical epidemiology; critical care; postoperative ventilation; quality measures; risk assessment

Introduction

Each year, 3–8% of patients develop postoperative pulmonary complications (PPC) amounting to approximately 2.5 million patients in the USA alone [1, 2]. Postoperative pulmonary complications are associated with an increased risk of mortality, morbidity and postoperative discharge to a nursing home [3–7]. Commonly considered respiratory complications after surgery include pneumonia, pulmonary oedema, atelectasis and acute respiratory failure, potentially resulting in tracheal re-intubation [8]. Unplanned tracheal re-intubation after surgery was included as a quality benchmark measure in the Quality and Resource Use reports by the Centers for Medicare and Medicaid Services [9].

To enhance the quality and safety of peri-operative patient care, the Agency for Healthcare Research and Quality recommends risk assessment for postoperative respiratory failure [10]. We developed previously the Score for Prediction of Postoperative Respiratory Complications (SPORC); this is a widely accepted, simple prediction model for development of PPCs with the primary endpoint of early (within 72 h of surgery) tracheal reintubation [11]. The SPORC has been advocated by the Agency for Healthcare Research and Quality and has been implemented in recommendations reflecting state of the art knowledge for pre-operative patient screening [10, 12].

Our primary aim was to develop and externally validate an improved prediction score (SPORC-2) that considered both pre- and intra-operative predictors, and which could be used by clinicians to assess patients for the risk of early postoperative tracheal re-intubation. Our secondary aim was to compare this new tool for stratified risk assessment to existing models that focus only on pre-operative predictors. We hypothesised that the addition of intra-operative variables to pre-operative risk assessment would significantly improve the predictive value of the model.

Methods

In this hospital registry study, patient data from two independent healthcare networks (Partners Healthcare and Beth Israel Deaconess Medical Center (BIDMC), Boston, MA, USA) were analysed. Both the Partners Institutional Review Board and the Committee on Clinical Investigations at Beth Israel Deaconess Medical Center approved this study and waived requirement for patient consent. This manuscript adheres to the applicable TRIPOD guidelines [13].

For score development, all patients undergoing non-cardiac surgery between 1 January 2007 and 31 December 2015 at Massachusetts General Hospital, Boston, and two community hospitals were screened for eligibility. For external validation, we considered all non-cardiac surgical patients at BIDMC between 1 October 2005 and 30 September 2017. All patients included in the study underwent general anaesthesia with tracheal intubation and post-procedural extubation in the operating theatre. For both cohorts, patients with any of the following criteria were not studied: ambulatory surgery; age < 18 years; ASA physical status 6; surgery within 10 days before the index procedure; and missing data-points of predictor variables. We accounted for missing data by applying multiple imputation by chained equations in both cohorts (see also Supporting Information, Section 7).

The primary outcome was early tracheal re-intubation after post-procedural extubation. This was defined as requirement for tracheal re-intubation within 72 h of surgery with subsequent mechanical ventilation in the operating theatre, recovery area or intensive care unit (ICU). Tracheal intubations within 72 h of the index procedure for an additional surgical procedure were not included in the primary outcome. In the development cohort, tracheal re-intubation was captured based on current procedural terminology codes for tracheal intubation and mechanical ventilation management. The primary outcome variable has been validated previously in the development cohort (see also Supporting Information, Section 1.1) [11]. In the validation cohort, tracheal re-intubation was captured based on time stamps for post-procedural tracheal extubation and mechanical ventilation and mechanical ventilation tracheal re-intubation was captured based on time stamps for post-procedural tracheal extubation and mechanical ventilation after surgery available from the respiratory therapists' database.

We considered both pre-operative (comorbidities and pre-operatively available measures of procedural risk) and intra-operative variables for the development of SPORC-2. A detailed definition of all predictors can be found in the Supporting Information (Section 3.1, Table S1). All five predictors of the previously reported SPORC were considered: ASA physical status 3; history of chronic pulmonary disease; history of heart failure; emergency surgery; and high-risk surgical services (vascular surgery; transplant surgery; neurosurgery; thoracic surgery; general surgery; and burn surgery) [11]. Furthermore, we considered surgical complexity as quantified by the procedural severity score [14]. Intra-operative candidate variables were selected a priori based on recent literature, biological plausibility and clinical reasoning and included: desaturation (S_pO_2 90%) 5 min after tracheal intubation [7]; duration of surgery; median fraction of inspired oxygen (F_IO_2) [15]; noradrenaline equivalent dose of vasopressors; dose of neuromuscular blocking agents (NMBA) (expressed as multiples of NMBA dose needed to reduce twitch height by 95%) [16, 17]; fluid volume [18]; oral morphine equivalent dose [19, 20]; total administered fentanyl dose;

intra-operative transfusion of packed red blood cells; use of volatile anaesthetic agents [21]; and the absence of lung-protective ventilation (defined as driving pressure [plateau pressure – positive end-expiratory pressure (PEEP)] > 15 mmHg) [22, 23].

The initial full multivariable logistic regression model contained all a priori defined pre- and intra-operative candidate predictors. To derive the final model, we used stepwise backward regression with p values < 0.05 to retain predictors from the set of candidate variables. This was followed by bootstrap resampling (n = 500 repetitions) to confirm predictor robustness and avoid overfitting. All predictors retained in the final model were assigned a weighting value by dividing the respective beta coefficient by the smallest beta coefficient among all final predictors. Weighting values were rounded to the nearest integer and multiplied by the respective predictor, thus estimating the final prediction score. All analyses were performed using Stata version 13 (Stata Corp LLC, College Station, TX, USA) or RStudio v1.0136 (RStudio. Inc, Boston, MA, USA).

Model discrimination was assessed by performing C-statistics. The Brier Score was calculated to examine model accuracy. We further assessed model calibration by creating a calibration plot. The Hosmer-Lemeshow test was performed with a p value > 0.05 indicating good model fit [24]. Analyses on model performance refer to the regression model with the prediction score as a single independent variable. The score variable represents the sum of all final predictors multiplied by their respective weighting values. In the independent validation cohort, we applied SPORC-2 to all patients and assessed model performance utilising C-statistics, Brier Score, and calibration plot. We compared the newly derived SPORC-2 with the original SPORC. Comparison of the predictive value of the two models was performed based on the respective C-statistics results. Furthermore, we performed net reclassification improvement analysis to assess whether the addition of unique procedure-related and intra-operative predictors to the original SPORC led to enhanced risk assessment.

Postoperative pulmonary complications may lead to increases in overall organ dysfunction, mortality and utilisation of healthcare resources [25]. As part of an exploratory analysis, we examined the association between SPORC-2 and further outcomes including: postoperative hospital duration of stay (defined as the number of hospitalised days after surgery); 30-day all-cause mortality; and 30-day re-admission rate. Analyses on the consequences of a high SPORC-2 were conducted in a dataset combining the development and validation cohorts. We compared the discriminatory ability of SPORC-2 with another existing model, the Assess Respiratory Risk in Surgical Patients in Catalonia (ARISCAT), applying the same comparison analyses as described above [26]. Results are described in the Supporting Information (Section 6.1.1). The pre-operative SPORC-2 consists of all pre-operatively available predictors (see also Supporting Information, Table S3). We compared the pre-operative SPORC-2 to the full SPORC-2. Results are described in the Supporting Information (Section 6.1.2).

Results

The final development cohort consisted of 90,893 patients, of whom 699 patients (0.8%) had their trachea re-intubated within 72 h of surgery (Table 1). At BIDMC, a total of 67,046 patients were studied; in the final validation cohort, 587 patients (0.9%) required tracheal re-intubation within 72 h of surgery (Fig. 1 and Supporting Information, Section 2).

A total of 12 predictors were retained in the final model for SPORC-2: five pre-operative variables (ASA physical status 3; heart failure; chronic pulmonary disease; emergency surgery; and high procedural severity score (32–48 and 49)) [14]; and seven intraoperative factors (desaturation (S_pO_2 90%) 5 min after tracheal intubation; duration of surgery > 140 min and > 225 min; high median F_IO_2 (> 0.61); high total noradrenaline equivalent dose (> 0.18 mg); intra-operative transfusion of packed red blood cells; the absence of volatile anaesthetic agents; and lack of lung-protective ventilation patterns). The final prediction score reached a maximum score point value of 46 points (Fig. 2). Predicted risks of postoperative tracheal re-intubation for increasing score point values derived from the development cohort are shown in the Supporting Information (Table S4).

In the development cohort, the final model had an area under the receiver operating characteristic curve (AUC) of 0.84 (95% CI 0.82–0.85) showing excellent discriminative ability (Fig. 3) [24]. The uncertainty of 0.01 has to be considered in relation to the relatively low incidence of the outcome. Predictions ranged from 0.2% to 60.3% and were well calibrated (reliability 0.002). The Hosmer–Lemeshow test was not significant (p = 0.06), indicating good model fit. In the validation cohort, C-statistics proved good discriminative ability (AUC 0.75 (95% CI 0.73–0.77)) (Fig. 3). The Brier Score of 0.012 (uncertainty 0.010, reliability 0.002) confirmed the results of the development cohort. Figure 4 shows the model calibration in the development and validation cohorts.

When comparing the newly derived SPORC-2 to SPORC, the AUCs showed significantly improved model discrimination in the development cohort (0.84 (95% CI 0.82–0.85) vs. 0.75 (95% CI 0.73–0.77), respectively; p < 0.001) (Fig. 5a). In addition, risk categorisation was significantly improved in net reclassification improvement (NRI) analyses (additive NRI 0.27; p < 0.001). Overall, 232 (33.1%) patients were adequately reclassified into a higher, and 16,209 (18.0%) patients into a lower risk category when considering the additional intra-operative predictors of SPORC-2 (see also Supporting Information, Section 5 and Table S2a). In the external validation cohort, SPORC-2 performed significantly better than SPORC (AUC 0.75 (95% CI 0.73–0.77 vs. 0.69 (95% CI 0.67–0.70), respectively; p < 0.001) (Fig. 5b). Net reclassification improvement results derived from the external validation cohort are described in the Supporting Information (Section 5, Table S2b).

In the exploratory analyses, when combining the development and validation cohorts, 1286 (0.8%) of 157,939 patients required early tracheal re-intubation after surgery, compared with 156,653 patients (99.2%) who did not. The requirement for early tracheal re-intubation was associated with increased 30-day mortality (11.0% vs. 0.7%, absolute risk difference 10.4% (95%CI 8.7–12.1%)), 30-day re-admission rates (15.8% vs. 8.3%, absolute risk difference

7.5% (95%CI 5.5–9.5%)) and postoperative hospital duration of stay (median (IQR [range]), 10 (6–16 [1–326]) days vs. 4 (2–6 [1–288]) days).

Discussion

We developed and externally validated SPORC-2, a prediction instrument for early tracheal re-intubation after surgery. The incorporation of intra-operative predictors resulted in a significantly improved and clinically more relevant prediction model, with enhanced discriminative ability and risk categorisation in both the development and validation cohorts.

The primary end-point of the present study was tracheal re-intubation within 72 h of surgery. Early postoperative tracheal re-intubation is one of the most rigorous markers for persistent respiratory failure [3, 27]. In our study, early tracheal re-intubation was associated with increased all-cause mortality within 30 days of surgery, 30-day re-admission rate and hospital duration of stay. The Agency for Healthcare Research and Quality's recommendation to assess the risk of early tracheal re-intubation underlines the need for evidence-based prediction instruments for PPCs. The inclusion of tracheal re-intubation as a quality benchmark in the Quality and Resource Use reports by the Centers for Medicare and Medicaid Services confirms its importance as an adverse patient-centred outcome [9].

Several predictors in the presented score relate to procedural risk independent of the individual intra-operative course, such as emergency versus elective surgery, procedural severity score and duration of surgery. In many academic centres, predicted duration of surgery data are available pre-operatively, and administrators use these data for optimal operating room scheduling [28]. Our prediction instrument includes several potentially modifiable intra-operative predictors, such as high median F_IO_2 , absence of volatile anaesthetic agents, high vasopressor dose, absence of lung-protective ventilation and intra-operative transfusion of packed red blood cells.

We found a strong association between high median $F_{I}O_2$ (> 0.61) and early tracheal reintubation after surgery, which is in line with previous findings [15]. A high $F_{I}O_2$ in general may be a marker for pulmonary dysfunction that would not be accounted for by other predictor variables. Adverse effects of higher inspiratory oxygen concentrations may be explained by increased absorption atelectasis [29], as well as increased production of reactive oxygen species with resultant oxidative stress injury to the lungs and other organ systems [30]. Atelectasis may contribute to additional pulmonary complications, thereby increasing the likelihood of requiring postoperative tracheal re-intubation [31]. The negative effects of high inspired oxygen concentrations need to be weighed against the clinical requirement of avoiding and safeguarding against potentially injurious hypoxaemia.

Our finding of the absence of volatile anaesthetic agents as a significant predictor for early tracheal re-intubation confirms previous results. Volatile anaesthetic agents are bronchodilators and immunomodulators [32], and their intra-operative use is dose-dependently associated with lower odds of PPCs [21].

We observed intra-operative haemodynamic instability, expressed as the requirement of high vasopressor doses and transfusion of packed red blood cells, to be associated with an

increased risk of early tracheal re-intubation. The predictive value of high vasopressor dose as a risk factor for the development of PPCs was shown previously in the trial by Neto et al. [33]. Both high vasopressor dose and intra-operative transfusion of packed red blood cells are indicators of haemodynamic instability due to ongoing intra-operative blood loss and/or a higher degree of distributive shock, conditions that are associated with an increased risk of PPCs.

Increasing and individualising PEEP levels, as well as decreasing plateau airway pressure, may have lung-protective effects [22, 34]. Accordingly, the absence of lung-protective ventilation patterns was associated with an increased risk of early tracheal re-intubation, which is in line with previous work [22, 35]. It is possible to modify intra-operative driving pressure by optimising flow characteristics during intra-operative mechanical ventilation [35].

Comparison of SPORC-2 with other published tools for pre-operative risk assessment (e.g. SPORC, ARISCAT) showed enhanced performance of SPORC-2 [11, 26]. One-third of patients in the score development cohort were adequately reclassified into higher risk categories when applying SPORC-2 instead of SPORC. To be optimally effective, the SPORC-2 tool should be used for stratified risk prediction. For example, SPORC-2 may be able to assign a patient score using data from pre-operative anaesthetic assessment that includes medical history (chronic pulmonary disease, heart failure), ASA physical status, emergency status and an automatically calculated procedural severity score. This information will be important for intra-operative resource allocation and postoperative critical care requirements. Intra-operatively, the clinician will integrate information provided by device data and documentation of events (e.g. lack of lung-protective ventilation, use of blood products, etc.) to generate a full SPORC-2 in real time. This information may change postoperative management; for example, ICU bed allocation depends on intra-operative risk assessment [36]. Given the expansion of electronic health records and anaesthesia information management systems, one would presume that generating SPORC-2 in real time using patient baseline characteristics, biomedical device and medical record data would be straightforward. It is encouraging to see a new breed of technologies with advanced medical device connectivity, powerful processing capability and real-time decision support enter the anaesthesia software market [37-39].

The utilisation of intra-operative predictors is an important distinctive feature of our prediction model. The majority of previous studies on the prediction of PPCs focused on predictors only available pre-operatively [26, 40, 41]. The trial by Neto et al, which derived a moderately performing prediction model comprising six pre-operative, five intra-operative and two procedure-related risk factors, was limited by its relatively small sample size and lack of external validation [33]. Further studies investigating intra-operative predictors of PPCs were similarly limited by small sample size and lack of external validation [2, 42]. In the present study, we externally validated our score in an independent cohort with a different case mix and outcome incidence. The utilisation of two different cohorts with high-quality, validated data allowed us to prove the generalisability of the newly developed model.

This study has several limitations. The observational nature of the study and use of administrative data and billing codes confer risks of unmeasured confounding factors and bias. Second, despite the granularity of the databases, the independence of the cohorts used for SPORC-2 development and validation, and the robustness of our findings, there are some limitations to the scope of the subjects captured. Regarding PPCs, risk assessment of subjects whose tracheas remained intubated at the end of the procedure would be of comparative interest. It remains unclear whether there are differences in characteristics and outcomes between patients whose tracheas remain intubated at the end of surgery compared with those whose tracheas are primarily extubated and subsequently re-intubated later.

In summary, we have derived and validated SPORC-2, an instrument for stratified assessment of a patient's risk of early tracheal re-intubation after surgery. This tool may improve peri-operative resource allocation for surgical patients and will help clinicians fulfil impending reporting requirements. It may also prove beneficial in supporting clinicians in their efforts to advance patient safety and improve overall quality of care.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

Acknowledgements

This work was funded by philanthropic donors Dr Jeffrey and Judy Buzen. The donors were not involved in the study design, data collection, data analysis, data interpretation or writing of the manuscript and supporting materials. HL is an employee at Talis Clinical LLC, a company that offers secure web-based online medical records. TH reports grants from NINDS and NIGMS, and personal fees from Headache, Anesthesiology and Cephalalgia, outside the submitted work. ME receives funding from philanthropic donors Dr Jeffrey and Judy Buzen. No other external funding or competing interests declared.

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Figure 1.

Study flow chart. Multiple exclusion criteria may apply.

Pre-operative variables					
Heart failure					
Emergency procedure					
ASA physical status ≥ 3					
PSS Morbidity ≥ 49					
PSS Morbidity 32–48					
Chronic pulmonary disease	_				
Intra-operative variables					
Duration of surgery > 225 min		-			
Duration of surgery > 140 min		_	-		
Post-tracheal intubation desaturation $\leq 90\%$					-
Median fraction of inspired oxygen > 0.61					
Administration of packed red blood cells					
Absence of lung-protective ventilation					
Noradrenaline equivalent > 0.18 mg			-		
Absence of volatile anaesthetics		-			
Score point values	0	1	2	3	4

Figure 2.

Factors used in the score for the prediction of early tracheal re-intubation after surgery (Score for Prediction of Postoperative Respiratory Complications-2). Score point values of applicable predictors are summed up to create total score. PSS, procedural severity score.

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Figure 3.

Model discrimination (C-statistics) for the score for the prediction of early tracheal reintubation after surgery (Score for Prediction of Postoperative Respiratory Complications-2) in development (Fig. 3a) and external validation (Fig. 3b) cohorts.



Figure 4.

Calibration plots for the score for the prediction of early tracheal re-intubation after surgery (Score for Prediction of Postoperative Respiratory Complications-2) in development (Fig. 4a) and external validation (Fig. 4b) cohorts.

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Figure 5.

Comparison of model discrimination (C-statistics) for Score for Prediction of Postoperative Respiratory Complications (SPORC-2) (dashed lines) and SPORC (solid lines) to predict early tracheal re-intubation after surgery in development (Fig. 5a) and validation (Fig. 5b) cohorts.

Table 1

Clinical characteristics of the development and validation cohort stratified by the presence or absence of early postoperative tracheal re-intubation. Values are number (proportion), mean (SD) or median (IQR [range]).

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	Development cohort (n = 90,893)		Validation cohort $(n = 67,046)$	
	No tracheal re-intubation (n = 90,194)	Tracheal re-intubation (n = 699)	No tracheal re-intubation (n = 66,459)	Tracheal re-intubation (n = 587)
Sex; male	40,523 (45%)	372 (53%)	28,050 (42%)	298 (51%)
Age; years	56 (17)	63 (16)	56 (17)	63 (15)
Body mass index; kg.m ⁻²	28.6 (7.1)	28.5 (7.8)	28.8 (7.3)	28.9 (7.9)
ASA physical status	2 (2–3 [1–5])	3 (2–3 [1–5])	2 (2–3 [1–5])	3 (2–3 [1–5])
Chronic pulmonary disease	16,943 (19%)	228 (33%)	12,353 (19%)	188 (32%)
Heart failure	7395 (8%)	240 (34%)	4219 (6%)	101 (17%)
Emergency surgery	4,711 (5%)	104 (15%)	4,965 (8%)	88 (15%)
High-risk surgical services ^a	40,348 (45%)	413 (59%)	30,267 (46%)	301 (51%)
Duration of procedure; min	177 (122–253 [9–1153])	232 (146–343 [31–885])	156 (112–222 [0–1147])	183 (124–267 [15–874]
Duration of intra-operative MAP< 55 mmHg; min	0 (0–2 [0–614])	2 (0-6 [0-258])	1 (0–3 [0–243])	1 (0-4 [0-59])
Procedural severity score for morbidity	41 (18)	54 (17)	39 (16)	49 (17)
Packed red blood cells; 0 units	86,299 (96%)	572 (82%)	64,250 (97%)	496 (85%)
Packed red blood cells; $1-2$ units	3189 (4%)	93 (13%)	1981 (3%)	76 (13%)
Packed red blood cells; 3 units	706 (1%)	34 (5%)	228 (0%)	15 (3%)
$Fluids$, 1^b	1.5 (1.0–2.5 [0.0–30.8])	2.0 (1.0–3.5 [0.0–20.2])	3.5 (2.5–5.0 [0.0–34.0])	4.0 (2.5–6.5 [0.0–24.9])
Oral morphine equivalent of total long-acting intra-operative opioid dose; mg	15 (2–27 [0–545]	7 (0–24 [0–408])	14 (0-24 [0-558])	14 (0–24 [0–102])
95% effective dose of total NMBA; mg	2.7 (1.7-4.1 [0.0-46.7])	3.3 (2.1–5.1 [0.0–19.0])	2.2 (1.2–3.2 [0.0–40.7])	2.6 (1.5-4.0 [0.0-17.8])
Neostigmine dose;mg	2 (0-4 [0-23])	3 (0-4 [0-10]	3 (0-4 [0-20])	3 (0-4[0-8])
Dose equivalent of inhalational anaesthetic; MAC	0.6 (0.4–0.8 [0.0–2.5])	0.5 (0.3–0.8 [0.0–1.6])	0.9 (0.7–1.1 [0.0–3.0])	0.9 (0.7–1.1 [0.0–2.4])
Noradrenaline equivalent of vasopressor dose; mg	0.1 (0.0–0.3 [0.0–97.5]	0.3 (0.1–0.8 [0.0–91.0]	0.0 (0.0-0.1 [0.0-72.5])	0.1 (0.0–0.5 [0.0–45.7])
S_pO_2 5 min post-tracheal intubation; %	100 (99–100 [45–100])	100 (99–100 [80–100])	99 (99–100 [60–100])	100 (98–100 [82–100]
F_1O_2 ; %	52 (38–58 [21–100])	57 (46–77 [21–100]	54 (48-60 [21-100])	58 (52–78 [21–100])

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	Development cohort $(n = 90, 893)$		Validation cohort $(n = 67,046)$	
	No tracheal re-intubation (n = 90,194)	Tracheal re-intubation (n = 699)	No tracheal re-intubation (n = 66,459)	Tracheal re-intubation $(n = 587)$
Lung-protective ventilation $^{\mathcal{C}}$	55,417 (61%)	354 (51%)	19,163 (29%)	148 (25%)

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ASA, American Society of Anesthesiologists; MAP, mean arterial pressure; NMBA, neuromuscular blocking agent; MAC, minimum alveolar concentration; FIO2, fraction of inspired oxygen; SpO2, peripheral capillary oxygen saturation.

 $^{a}_{1}$ Includes vascular surgery, transplant surgery, neurosurgery, thoracic surgery, general surgery and burns.

bCrystalloid:colloid ratio 1:1.5.

c²Lung-protective ventilation was defined as driving pressure 15 mmHg(median plateau pressure-median positive end-expiratory pressure).