# Predictors of mortality and cost among surgical patients requiring rapid response team activation

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**Background:** Prior studies of rapid response team (RRT) implementation for surgical patients have demonstrated mixed results with respect to reductions in poor outcomes. The aim of this study was to identify predictors of in-hospital mortality and hospital costs among surgical inpatients requiring RRT activation.

**Methods:** We analyzed data prospectively collected from May 2012 to May 2016 at The Ottawa Hospital. We included patients who were at least 18 years of age, who were admitted to hospital, who received either preoperative or postoperative care, and and who required RRT activation. We created a multivariable logistic regression model to describe mortality predictors and a multivariable generalized linear model to describe cost predictors.

**Results:** We included 1507 patients. The in-hospital mortality rate was 15.9%. The patient-related factors most strongly associated with mortality included an Elixhauser Comorbidity Index score of 20 or higher (odds ratio [OR] 3.60, 95% confidence interval [CI] 1.96–6.60) and care designations excluding admission to the intensive care unit and cardiopulmonary resuscitation (OR 3.52, 95% CI 2.25–5.52). The strongest surgical predictors included neurosurgical admission (OR 2.09, 95% CI 1.17–3.75), emergent surgery (OR 2.04, 95% CI 1.37–3.03) and occurrence of 2 or more operations (OR 1.73, 95% CI 1.21–2.46). Among RRT factors, occurrence of 2 or more RRT assessments (OR 2.01, 95% CI 1.44–2.80) conferred the highest mortality. Increased cost was strongly associated with admitting service, multiple surgeries, multiple RRT assessments and medical comorbidity.

**Conclusion:** RRT activation among surgical inpatients identifies a population at high risk of death. We identified several predictors of mortality and cost, which represent opportunities for future quality improvement and patient safety initiatives.

**Contexte** : Les études sur la mobilisation d'équipes d'intervention rapide (EIR) auprès de patients en chirurgie ont donné des résultats mitigés quant à la réduction des issues négatives. La présente étude visait à déterminer les facteurs prédictifs de coûts pour les hôpitaux et de mortalité chez les patients en chirurgie nécessitant la mobilisation d'une EIR.

**Méthodes** : Nous avons analysé des données recueillies de manière prospective de mai 2012 à mai 2016 à l'Hôpital d'Ottawa. Nous avons inclus les patients hospitalisés de 18 ans et plus qui ont reçu des soins préopératoires ou postopératoires et qui ont nécessité l'intervention d'une EIR. Nous avons ensuite créé un modèle de régression logistique multivariée pour décrire les facteurs prédictifs de mortalité et un modèle linéaire généralisé multivarié pour décrire les facteurs prédictifs de coûts.

**Résultats** : Nous avons retenus 1507 patients. Le taux global de mortalité à l'hôpital était de 15,9 %. Les principaux facteurs de mortalité liés au patient étaient un indice de comorbidité d'Elixhauser supérieur ou égal à 20 (rapport de cotes [RC] 3,60, intervalle de confiance [IC] à 95 % 1,96–6,60) et des objectifs de soins excluant l'admission à l'unité des soins intensifs et la réanimation cardiorespiratoire (RC 3,52, IC à 95 % 2,25–5,52). Les principaux facteurs prédictifs liés aux interventions sont l'admission en neurochirurgie (RC 2,09, IC à 95 % 1,17–3,75), l'intervention chirurgicale d'urgence (RC 2,04, IC à 95 % 1,37–3,03) et le fait d'avoir subi au moins 2 opérations (RC 1,73, IC à 95 % 1,21–2,46). Parmi les facteurs liés aux EIR, la tenue d'au moins 2 évaluations par l'EIR s'accompagnait du mortalité le plus élevé (RC 2,01, IC à 95 % 1,44–2,80). L'augmentation des

coûts était étroitement associée au service d'admission, aux interventions chirurgicales multiples, aux évaluations multiples par l'EIR et à la comorbidité médicale.

**Conclusion :** La mobilisation d'EIR auprès de patients en chirurgie permet de mettre en évidence une population à risque élevé de décès. Nous avons découvert plusieurs facteurs prédictifs de mortalité et de coûts, dont on pourra se servir pour améliorer la qualité des soins et la sécurité des patients.

any hospitals have adopted rapid response teams (RRTs), often comprising multidisciplinary critical care providers including nurses, respiratory therapists and physicians, to provide urgent critical care expertise when activated.<sup>1</sup> The aim of these teams is to intervene when patients admitted to hospital have abnormal vital signs and biochemical abnormalities in the hope of preventing further deterioration.<sup>2</sup> Reasons for RRT activations typically include hypotension, respiratory distress, arrhythmias, altered level of consciousness, or concern of the clinical ward team of impending deterioration.

A recent systematic review evaluating RRTs concluded that implementation of RRTs was associated with a reduction in cardiac arrests among medical ward patients.<sup>3</sup> However, prior studies of RRT implementation for surgical patients have demonstrated mixed results with regard to reduction of critical events such as cardiac arrests, intensive care unit (ICU) admissions and in-hospital mortality.<sup>4-7</sup> Potential reasons for this variability between medical and surgical patients include attitude differences among physicians, attitude differences among nurses and fundamental differences in disease pathophysiology.<sup>5-10</sup>

Despite these critical differences, studies specifically examining the surgical population are limited. When institutional protocols for activation are inefficient or not implemented with input from front-line staff, studies have demonstrated noncompliance rates surpassing 75% even when activation criteria are met, and median times to activation as high as 16 hours.<sup>11</sup> Interventions tailored to the individual needs of each institution have been shown to be more effective in terms of promoting safety culture and improving patient outcomes.<sup>12,13</sup> To identify the most vulnerable patients in this surgical subpopulation who may potentially benefit from tailored RRT care and follow-up, we sought to identify specific predictors of poor outcome. The objective of this study was to identify the predictors of both in-hospital mortality and hospital costs among surgical inpatients requiring RRT activation.

#### METHODS

#### Cohort

This is a retrospective analysis of a prospectively collected patient cohort from 2 hospitals in The Ottawa Hospital network, a tertiary care hospital network in Ottawa, Canada. We included all patients who were admitted to the hospitals between May 2012 and May 2016 and met the following inclusion criteria: aged 18 years or older, admitted to a surgical service at either of the study sites, required RRT activation at any time during the index hospital admission, and received either preoperative or postoperative care. Patients were included regardless of their level of care designation. The 3 levels are as follows: active medical treatment including ICU admission and cardiopulmonary resuscitation (CPR), active medical treatment including ICU admission without CPR, and active medical treatment excluding ICU admission or CPR. Patients with cardiac arrests were excluded as the institutional "code blue" response team responds to such patients. In the event of an RRT activation that escalates to a code blue for cardiac arrest, the patient is tended to by both the RRT and the code blue response team; such patients were included in this cohort. Ethics approval was obtained from the Ottawa Health Science Network Research Ethics Board.

#### Setting and data collection

The Ottawa Hospital is a 1163-bed health care network that handles 160000 emergency visits, 50000 hospital admissions, 2500 ICU admissions and 35000 surgical cases per year. The hospital network serves as the regional referral centre for oncology, vascular surgery, neurosurgery and thoracic surgery. Each campus has a tertiarylevel, mixed medical–surgical ICU with a 28-bed capacity. The campuses use the same RRT activation criteria and institutional database although each caters to a different patient population mix.

The RRT at the Ottawa Hospital comprises a critical care physician, a critical care registered nurse with at least 5 years of critical care experience, and a respiratory therapist; the team can be activated to provide urgent medical care at the bedside. Nighttime RRT activations are managed by a resident physician on the critical care service with support from the registered nurse and respiratory therapist. Institutional criteria for RRT activation, presented in Appendix 1 (available at canjsurg.ca/017319-a1), include airway concerns, decreased level of consciousness, tachypnea, hypoxia, hypotension, arrhythmias and general worry of impending deterioration.<sup>12</sup> Health care providers are also encouraged to activate the RRT if they have concerns that are not outlined in the institutional criteria. Data for the RRT database are prospectively gathered by

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the responding registered nurse using a standardized data collection form at the time of patient assessment.

We linked prospectively collected data sets from the ICU RRT database with the institutional database, termed the Ottawa Hospital Data Warehouse (OHDW). The OHDW is a central repository of clinical and health administrative data that includes the patient registration system, clinical data repository, case-costing system and patient discharge abstracts. Linkage is performed using unique anonymized patient-level identifiers. The Ottawa Hospital's health administrative database has been widely used in previous health services research.<sup>1,13–15</sup> Data quality assessments were performed during development and are executed routinely as new data are added to ensure completeness and accuracy.

#### Predictors and outcome variables

Predictors of interest were selected on the basis of their clinical importance as guided by clinical expert opinion and the published literature. These included patient demographic characteristics, such as age and sex; medical comorbidity, defined using the Elixhauser Comorbidity Index;<sup>16</sup> medical fitness at the time of surgery, defined using the American Society of Anesthesiologists (ASA) classification system;<sup>17</sup> surgical characteristics, such as admitting service, occurrence of elective or emergent surgery and occurrence of multiple operations; and RRT activation characteristics, such as reason for activation (grouped by organ system) and occurrence of multiple activations. The Elixhauser Comorbidity Index score was extracted from the institutional database; it was calculated on the basis of the International Statistical Classification of Diseases and Related Health Problems, 10th revision diagnosis codes in administrative data and has been previously validated for the prediction of hospital costs and in-hospital mortality.<sup>16</sup> Previous work has demonstrated its superior discriminative ability over the Charlson Comorbidity Index score or modified Frailty Index score for the prediction of adverse events after orthopedic surgery.<sup>18</sup> The ASA class and surgical procedure data were extracted directly from the operative record of the first operation.

Data points for the RRT database are collected at the time of RRT assessment by the responding critical care nurse, while data points for the health administrative database are entered at the time of admission and discharge by trained clerical staff. For continuous variables, we investigated their functional forms and assessed for potential transformation with restricted cubic splines, fit with 3 knots. If the transformation improved the model fit, we categorized the continuous variable at the spline infection points. This minimizes inappropriate assumptions of linearity and arbitrary categorization thresholds that result in data waste and statistical inefficiency.<sup>19</sup> The remaining predictors were treated as categorical variables.

The dependent variables of interest were in-hospital mortality and total hospital admission costs. Total costs were determined from indirect costs (overhead operational fees) and direct costs (salaries and materials not including physician remuneration). The Ottawa Hospital uses standardized case-costing methodology developed by the Ontario Case Costing Initiative<sup>20</sup> and based on the Canadian Institute for Health Information Management Information System guidelines.<sup>14</sup> This case-costing algorithm, used in previous health services literature,<sup>14,15,21</sup> calculates total hospital costs on a hospital-specific basis using standardized methods and includes both direct and indirect costs standardized to 2018 Canadian dollars.

### Statistical analysis

For descriptive statistics, categorical variables are presented as counts and percentages, and continuous variables are presented as medians with interquartile ranges (IQRs) to account for nonnormal distributions. Variables with normal distributions are presented as means with standard deviations (SDs). To identify predictors of inhospital mortality, a multivariable logistic regression model was created on the basis of a priori selection of clinically important variables, including patient demographics, medical comorbidity, surgical characteristics and RRT activation characteristics, with associations presented as unadjusted and adjusted odds ratios (ORs) with 95% confidence intervals (CIs). We adhered to an event per variable ratio greater than 10,<sup>22</sup> and we intentionally avoided selection by univariate testing of association to minimize potential overfitting.<sup>19</sup> We used the C-statistic to characterize goodness of fit and corrected for optimism using 1000 bootstrap resamples for internal validation. Unadjusted associations are presented separately in Appendix 1. We used a multivariable generalized linear model (GLM) with a  $\gamma$  distribution to describe predictors of hospital costs, as this method accounts for highly skewed data without the need for transformation.<sup>23-25</sup> We similarly used a priori selection of clinically important variables and present associations as adjusted cost ratios (CRs). All statistical analyses were performed using SAS version 9.4 (SAS Institute) and SPSS version 24.0 (IBM) software. We used p < 0.05 as the threshold for statistical significance for all analyses.

#### RESULTS

Between May 2012 and May 2016, 1507 adult surgical inpatients met our eligibility criteria. The characteristics of the cohort are presented in Table 1. The median age was 71 (IQR 60–81) years, and 50.9% of patients were male. The median Elixhauser Comorbidity Index score was 4 (IQR 0–10), and the median ASA score was 3 (IQR 3–4). The most common admitting services were general surgery (23.2%) and orthopedics (29.0%). There were 898 (59.6%) surgeries classified as emergent, and 300 (19.9%) patients underwent 2 or more operations. The most common reasons for RRT activation were arrhythmias (23.0%), respiratory distress (18.0%) and decreased level of consciousness (17.3%).

Patient outcomes are presented in Table 2. There were 318 (21.1%) patients requiring ICU admission, and the overall in-hospital mortality rate was 15.9%. Among patients requiring RRT activation and ICU admission, the median hospital length of stay was 22.5 (IQR 11–41) days.

Characteristic	No. (%) of patients* n = 1507
Age, yr, median (IQR)	71 (60–81)
Sex, male	767 (50.9)
ICU admission within past yr	73 (4.8)
Admitting service	
General surgery	350 (23.2)
Orthopedics	437 (29.0)
Neurosurgery	158 (10.5)
Thoracic surgery	197 (8.6)
Trauma	19 (8.6)
Vascular surgery	199 (13.2)
Other	215 (14.3)
Level of care designation	
Full care including ICU admission and CPR	1164 (77.2)
Full care including ICU admission with no CPR	83 (5.5)
Full care excluding ICU admission	157 (10.4)
Reason for activation	
Airway concerns	41 (2.7)
Respiratory	271 (18.0)
Arrhythmias	346 (23.0)
Decreased level of consciousness	260 (17.3)
Hypotension	231 (15.3)
Other	314 (20.8)
Multiple RRT activations	358 (23.8)
Emergent (nonelective) operation	898 (59.6)
Multiple operations	300 (19.9)
Elixhauser Comorbidity Index score, median (IQR)	4 (0-0)
ASA class, median (IQR)	3 (3–4)

\*Unless indicated otherwise

Table 2. Clinical outcomes	
Outcome	No. (%) of patients* n = 1507
Need for ICU admission	318 (21.1)
Hospital length of stay, d, median (IQR)	
For entire cohort	15 (8–31)
For cohort requiring ICU admission	22.5 (11–41)
In-hospital mortality	240 (15.9)
ICU = intensive care unit; IQR = interquartile range. *Unless indicated otherwise.	

The results of the multivariable regression for predictors of in-hospital mortality among surgical patients requiring RRT activation are presented in Table 3. The patient factors most strongly associated with increased inhospital mortality included an Elixhauser Comorbidity Index score of 1–19 (OR 1.89, 95% CI 1.28–2.77), an Elixhauser Comorbidity Index score of 20 or higher (OR 3.60, 95% CI 1.96-6.60), increasing ASA class (OR 1.39, 95% CI 1.07–1.80) and designation of care excluding ICU admission and CPR (OR 3.52, 95% CI 2.25-5.52). The strongest surgical predictors included admission to a neurosurgical service (OR 2.09, 95% CI 1.17-3.75), occurrence of emergent surgery (OR 2.04, 95% CI 1.37-3.03) and occurrence of 2 or more operations (OR 1.73, 95% CI 1.21–2.46). Among RRT factors, occurrence of 2 or more RRT assessments (OR 2.01, 95% CI 1.44-2.80) or activation for respiratory distress (OR 1.95, 95% CI 1.24-3.07) conferred the highest associated odds of mortality. The naïve C-statistic was 0.78 and remained as such when

Table 3. Predictors of mortality	
Variable	Adjusted odds ratio (95% Cl)
Age ≥ 65 yr	1.35 (0.94–1.95)
Sex, male	1.00 (0.73–1.36)
ICU admission within past yr	1.05 (0.55–2.01)
Admitting service	
General surgery	1.44 (0.86-2.41)
Orthopedics	0.94 (0.54–1.61)
Neurosurgery	2.09 (1.17–3.75)
Trauma	1.62 (0.45–5.76)
Vascular surgery	1.71 (0.98–2.98)
Other	Ref.
Level of care designation	
Full care including ICU admission and CPR	Ref.
Full care including ICU admission with no CPR	1.70 (0.95–3.03)
Full care excluding ICU admission and CPR	3.52 (2.25–5.52)
Reason for activation	
Airway concerns	2.08 (0.90-4.84)
Respiratory	1.95 (1.24–3.07)
Arrhythmias	0.60 (0.36-1.00)
Decreased level of consciousness	1.58 (0.99–2.52)
Hypotension	1.28 (0.77–2.14)
Other	Ref.
Multiple RRT activations	2.01 (1.44–2.80)
Emergent (nonelective) operation	2.04 (1.37–3.03)
Multiple operations	1.73 (1.21–2.46)
ASA class (each increase in class)	1.39 (1.07–1.80)
ASA class 1	Ref.
Elixhauser Comorbidity Index score	
≤ 0	Ref.
1–19	1.89 (1.28–2.77)
≥20	3.60 (1.96–6.60)
ASA - American Society of Anesthesiologists: CI - confider	an interval: CDP -

ASA = American Society of Anesthesiologists; CI = confidence interval; CPR = cardiopulmonary resuscitation; ICU = intensive care unit; Ref. = reference; RRT = rapid response team.

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corrected for optimism with bootstrap validation over 1000 repeated samples.

The total hospital costs and cost allocation breakdown are presented in Table 4, indexed to 2018 Canadian dollars. The median total cost was \$30051 (IQR \$17496-\$56274), primarily driven by \$22440 (IQR \$13171-\$41890) in direct costs. The top 10% of high resource users had a mean cost of \$199567 (SD \$106380). The total cost for the cohort was \$76484573 and the cost per survivor was \$60367 for 1267 survivors. Nursing costs and operating room costs accounted for most of the total hospital costs.

The results of GLM modelling of costs are presented in Table 5. Patient factors most strongly associated with increasing cost included male sex (CR 1.10, 95% CI 1.02–1.18), an Elixhauser Comorbidity Index score of 20 or higher (CR 1.39, 95% CI 1.19-1.62) and increasing ASA class. In particular, an ASA class 5 level of preoperative medical fitness conveyed a 3.55-fold increase (95%) CI 2.26-5.60) in expected hospital admission costs. Patients with a level of care designation excluding ICU admission and CPR had an associated reduction in total costs (CR 0.84, 95% CI 0.75-0.95). Among admitting services, trauma team care was associated with higher cost (CR 2.19, 95% CI 1.57-3.05). The occurrence of multiple operations was associated with a 2.44-fold increase (95% CI 2.23-2.67) in costs. Among RRT factors, occurrence of 2 or more RRT assessments (CR 1.54, 95% CI 1.42-1.68) and activation for respiratory distress (CR 1.29, 95% CI 1.16-1.44) conferred the highest associated increase in costs.

Costs	Monetary value, Can\$
Costs, median (IQR)	
Direct costs	22440 (13171–41890)
Indirect costs	7614 (4420–14327)
Total costs	30051 (17 496–56274)
Costs for the top 10% of high-resource users, mean (SD)	199567 (106380)
Total costs for the entire cohort	76484573
Costs per survivor*	60367
Cost allocation, median (IQR)	
Food	770 (391–1561)
Health professionals	1619 (640–4215)
Laboratory investigations	1556 (735–3148)
Medical imaging	946 (377–2216)
Nursing	15–552 (7399–32–897)
Operating room	4423 (2918–7508)
Postanesthetic care unit	775 (459–1298)
Pharmacy	1742 (891–3851)
Respiratory therapists	575 (209–2133)

Variable Age ≥ 65 yr 0.93 (0.86-1.01) 1.10 (1.02-1.18) Sex, male 1.16 (0.98-1.37) ICU admission within past yr Admitting service General surgery 1.30 (1.15-1.46) 1.19 (1.06-1.35) Orthopedics 1.83 (1.58-2.13) Neurosurgery 2.19 (1.57-3.05) Trauma Vascular surgery 1.17 (1.02-1.35) Other Level of care designation Full care including ICU admission and CPR 1.39 (1.20-1.63) Full care including ICU admission with no CPR Full care excluding ICU admission and CPR 0.84 (0.75-0.95) Reason for activation Airway concerns 1.21 (0.96-1.52) Respiratory 1.29 (1.16-1.44) 0.97 (0.87-1.07) Arrhythmias Decreased level of consciousness 1 02 (0 91-1 14) Hypotension 0.98 (0.88-1.10) Other Multiple RRT activations 1.54 (1.42-1.68) 0.95 (0.87-1.03) Emergent (nonelective) operation Multiple operations 2.44 (2.23-2.67) ASA class 5 3.55 (2.26-5.60) 4 3.15 (2.08-4.77) 3 2.47 (1.64-3.73) 2 1

Table 5. Predictors of hospital admission cost

Adjusted cost ratio

(95% CI)

Ref.

Ref.

Ref

1.81 (1.18-2.79) Ref. Elixhauser Comorbidity Index score ≤ 0 Ref 1-19 1.06 (0.97-1.15) > 20 1.39 (1.19-1.62) ASA = American Society of Anesthesiologists; CI = confidence interval; CPR = cardiopulmonary resuscitation; ICU = intensive care unit; Ref. = reference; RRT = rapid response team.

### DISCUSSION

In this observational study of surgical inpatients requiring RRT assistance, we examined predictors of mortality and hospital admission costs. In this vulnerable population, approximately 1 in 6 patients died in hospital. The variables associated with the highest mortality included patient factors (increasing ASA class, higher Elixhauser Comorbidity Index score, level of care designation excluding ICU admission and CPR), surgical factors (admission to neurosurgical service, occurrence of emergent operation, occurrence of multiple operations) and RRT activation factors (activation for respiratory concerns and occurrence of multiple activations). Important predictors of hospital cost included admitting surgical service, occurrence of multiple surgeries, occurrence of multiple RRT

assessments, increasing ASA class and higher Elixhauser Comorbidity Index score.

These findings reflect a vulnerable population with high rates of morbidity and mortality and are consistent with the results of previous studies demonstrating increased odds of death and adverse events among postoperative patients with high ASA scores,<sup>26</sup> as well as among all patients (including nonsurgical patients) with recurrent deteriorations.<sup>27</sup> The neurosurgical population had the highest adjusted mortality among patients for all the admitting surgical subspecialties; these findings are consistent with this patient group's previously demonstrated mortality and cost burden.<sup>28</sup> Lastly, the high mortality rate associated with patients designated for care levels excluding ICU admission and CPR suggests that the priorities of RRT activation for this group should include improving symptom management and clarifying futility of life support, both of which are highly valuable.

With regard to resource utilization, we found that approximately 75% of total cost was driven by direct costs, naturally related to high patient-specific care needs, such as high-acuity nursing and operating room costs. Repeated utilization of RRT team assessments and surgical team resources was associated with higher costs, as expected. Similarly, a level of care designation precluding aggressive interventions, such as ICU admission or CPR, naturally reduced total costs. The importance of patient factors is particularly noteworthy. This is illustrated by the striking impact of poor preoperative medical fitness (ASA class 5) and high comorbidity burden (Elixhauser Comorbidity Index score  $\geq$  20), independent of age.

Although postoperative adverse events (AEs) are known to be major drivers of morbidity and hospital admission costs in the surgical patient population,15,29 there is little actual variability in incidence between the hospitals with the highest ("worst") and lowest ("best") AE rates.<sup>30</sup> However, the mortality rates for these AE rates, otherwise known as failure to rescue, at the worst hospitals were often 2 times higher than the rates at the best hospitals.<sup>31</sup> This phenomenon describes institutional differences in safety culture and quality of response to AEs; i.e., the ability to halt the potential transition from initial serious AE to eventually substantial morbidity and death. The RRT plays a crucial role in the "rescue" of these complex postoperative patients with acute deterioration by means of early identification, intervention and potential triage to higher-acuity care.<sup>32</sup> Similarly, some institutions use surgical step-down units (SDUs) to provide an intermediate level of care between the ICU and the general ward for patients at higher risk, including higher nurse-to-patient ratios or staffing by nurses with advanced training. There is evidence suggesting that SDUs provide a safe postoperative setting for such patients, resulting in fewer complications, improved pain control and reduced need for full intensive care.33

Previously validated approaches to patient safety improvement, notably the Comprehensive Unit-based Safety Program, emphasize identifying patient safety mistakes, using evidence-based interventions, placing value on the wisdom of front-line staff and empowering those staff to become actively involved in safety initiatives.<sup>34-36</sup> Similarly, the specific RRT framework is known to vary widely by institution and is often determined by local needs and resources.<sup>3</sup> This study therefore serves to identify several high-risk factors for mortality and increased cost among surgical patients requiring RRT activation and highlights a group of particularly vulnerable, high-cost patients. Although most of the variables identified are not directly modifiable, they present an opportunity to inform institutional protocols targeting high-risk patients who may benefit from earlier RRT involvement and closer followup care.

Although our cost analysis highlights predictors of high cost among RRT users, it does not specifically allow for an evaluation of resource utilization in an RRT-based care model compared with a non-RRT-based care model. In other words, the cost savings associated with the reduction of ICU utilization and unanticipated mortality remains unclear. These challenges are similarly faced by the existing literature evaluating the effectiveness of RRT care models, which is based primarily on nonrandomized and observational studies, focusing on clinical outcomes.<sup>32</sup>

Currently, our institution advocates for and supports liberal use of RRT activation, including dedicated foundational teaching to surgical residents regarding their reasons for activation and potential benefits. In addition, all residents completing critical care rotations will participate in the overnight RRT with oversight from an attending physician. However, we have yet to implement mandatory institutional interventions (such as ICU admission or palliative care discussion) aside from at least 1 follow-up assessment by an attending intensivist.

Future opportunities for further research and understanding of actionable items in this vulnerable population should focus on implementation factors of RRTs, including differences between physician- and nurse-initiated activations and the impact of delayed activation.

Our study has several strengths, including systematic record linkage across a number of high-quality data sets from a multihospital health care network, robust and modern methodologic approaches for prediction modelling and validation, and granular resource utilization data.

#### Limitations

Limitations of this study relate primarily to the observational nature of the data set. The identification of several factors highly associated with increased mortality and total hospital costs does not necessarily imply causation or even that modifying such factors would result in improved

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outcomes. Although our institution does not use formal SDUs, there are slight variations in nurse-to-patient ratios and nursing protocols between and within surgical wards, which we were unable to account for. Although previous literature has noted fewer ICU admissions and decreased length of stay with SDU implementation,<sup>33</sup> to our knowledge there is no existing evidence to characterize the association between SDU implementation and RRT utilization.

In addition, we were unable to comment on the cases of RRT activation noncompliance where activation criteria were met but did not result in RRT activation and assessment. Instead, these findings should serve as grounds for hypothesis generation and should prompt institutions to reflect on the specific vulnerabilities of their own RRT system when dealing with surgical patients.

Despite robust attempts to minimize confounding and overfitting, we were unable to control for all factors, including the degree of physiologic derangement as well as day–night differences, especially given the difference in training between the attending intensivist during the day and the resident physician at night.

Although 2 large hospitals contributed to this patient cohort, they exist within a single health care network and are influenced by similar local ICU and surgical practices that may limit generalizability. In addition, while each hospital uses the same RRT activation criteria and institutional database, we were unable to account for the specific patient population mix at each site with separate analyses. Lastly, we were unable to capture data regarding the nature and timing of interventions provided by the RRT, including resuscitation and medications.

### CONCLUSION

We found that RRT activation among surgical inpatients highlights a particularly vulnerable population at high risk of mortality and should lead to a higher level of scrutiny and diligence. We described several predictors of increased mortality and high resource utilization that may serve to more efficiently identify and tailor RRT care and follow-up as part of future patient safety initiatives.

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#### References

- 1. Fernando SM, Reardon PM, Bagshaw SM, et al. Impact of nighttime rapid response team activation on outcomes of hospitalized patients with acute deterioration. *Crit Care* 2018;22:67.
- 2. Davies O, DeVita MA, Ayinla R, et al. Barriers to activation of the rapid response system. *Resuscitation* 2014;85:1557-61.
- Winters BD, Weaver SJ, Pfoh ER, et al. Rapid-response systems as a patient safety strategy. *Ann Intern Med* 2013;158:417-25.
- Sarani B, Palilonis E, Sonnad S, et al. Clinical emergencies and outcomes in patients admitted to a surgical versus medical service. *Resuscitation* 2011;82:415-8.
- Moroseos T, Bidwell K, Rui L, et al. Rapid response team implementation on a burn surgery/acute care ward. *J Burn Care Res* 2014; 35:21-7.
- Jones D, Bates S, Warrillow S, et al. Effect of an education programme on the utilization of a medical emergency in a teaching hospital. *Intern Med J* 2006;36:231-6.
- Jones D, Opdam H, Egi Moritoki E, et al. Long-term effect of a medical emergency team on mortality in a teaching hospital. *Resuscitation* 2007;74:235-41.
- Sarani B, Sonnad S, Bergey M, et al. Resident and RN perceptions of the impact of a medical emergency team on education and patient safety in an academic center. *Crit Care Med* 2009;37:3091-6.
- Baxter AD, Cardinal D, Hooper J, et al. Medical emergency teams at the Ottawa Hospital: the first two years. *Can J Anaesth* 2008;55:223.
- Oake N, Taljaard M, van Walraven C, et al. The effect of hospital acquired *Clostridium difficile* infection on in-hospital mortality. *Arch Intern Med* 2010;170:1804-10.
- 11. Downey AW, Quach JL, Haase M, et al. Characteristics and outcomes of patients receiving medical emergency team review for acute change in conscious state or arrhythmias. *Crit Care Med* 2008;36:477-81.
- Vigorito MC, McNicoll L, Adams L, et al. Improving safety culture results in Rhode Island ICUs: lessons learned from the development of action-oriented plans. *Jt Comm J Qual Patient Saf* 2011;37:509-14.
- Khan RM, Ajuaid M, Ageel H, et al. Introducing the Comprehensive Unit-Based Safety Program for mechanically ventilated patients in Saudi Arabian Intensive Care Units. *Ann Thorac Med* 2017;12:11-6.

- Ronksley PE, McKay JA, Kobewka DM, et al. Patterns of health care use in a high-cost inpatient population in Ottawa, Ontario: a retrospective observational study. *CMAJ Open* 2015;3:E111-8.
- Reardon PM, Fernando SM, van Katwyk S, et al. Characteristics, outcomes and cost patterns of high-cost patients in the intensive care unit. *Crit Care Res Pract* 2018; doi: 10.1155/2018/5452683.
- Elixhauser A, Steiner C, Harris DR, et al. Comorbidity measures for use with administrative data. *Med Care* 1998;36:8-27.
- 17. Daabiss M. American Society of Anaesthesiologists physical status classification. *Indian J Anaesth* 2011;55:111-5.
- Ondeck NT, Bovonratwet P, Ibe IK, et al. Discriminative ability for adverse outcomes after surgical management of hip fractures: a comparison of the Charlson Comorbidity Index, Elixhauser Comorbidity Measure, and Modified Frailty Index. *J Orthop Trauma* 2018;32:231-7.
- Steyerberg EW. Clinical prediction models: a practical approach to development, validation and updating. Cham (Switzerland): Springer; 2009.
- Ontario Ministry of Health and Long-Term Care. Ontario Case Costing Initiative. Toronto: Ontario Ministry of Health and Long-Term Care; 2017. Available: www.ontario.ca/data/ontario-case -costing-initiative-occi/ (accessed 2019 Oct. 15).
- McIsaac DI, Abdulla K, Yang H, et al. Association of delay of urgent or emergency surgery with mortality and use of health care resources: a propensity score–matched observational cohort study. *CMAJ* 2017;189:E905-12.
- Peduzzi P, Concato J, Kemper E, et al. A simulation study of the number of events per variable in logistic regression analysis. *J Clin Epidemiol* 1996;49:1373-9.
- Moran JL, Solomon PJ, Peisach AR, et al. New models for old questions: generalized linear models for cost prediction. *J Eval Clin Pract* 2007;13:381-9.
- Mihaylova B, Briggs A, O'Hagan A, et al. Review of statistical methods for analysing healthcare resources and costs. *Health Econ* 2011; 20:897-916.
- Austin PC, Ghali WA, Tu JV. A comparison of several regression models for analysing cost of CABG surgery. *Stat Med* 2003;22:2799-815.

- 26. Hopkins TJ, Raghunathan K, Barbeito A, et al. Associations between ASA physical status and postoperative mortality at 48 h: a contemporary dataset analysis compared to a historical cohort. *Perioper Med* (*Lond*) 2016;5:29.
- Fernando SM, Reardon PM, Scales DC, et al. Prevalence, risk factors and clinical consequences of recurrent activation of a rapid response team: a multicenter observational study. *J Intensive Care Med* 2019;34:782-9.
- Fernando SM, Reardon PM, Dowlatshahi D, et al. Outcomes and costs of patients admitted to ICU due to spontaneous intracranial hemorrhage. *Crit Care Med* 2018;46:e395-403.
- Dimick JB, Chen SL, Taheri PA, et al. Hospital costs associated with surgical complications: a report from the private sector national surgical quality improvement program. *J Am Coll Surg* 2004;199:531-7.
- Ghaferi AA, Birkmeyer JD, Dimick JB. Variation in hospital mortality associated with inpatient surgery. N Engl J Med 2009; 361:1368-75.
- Ghaferi AA, Dimick JB. Understanding failure to rescue and improving safety culture. *Ann Surg* 2015;261:839-40.
- Barocas DA, Kulhalli CS, Ehrenfeld JM, et al. Benchmarking the use of a rapid response team by surgical services at a tertiary care hospital. *J Am Coll Surg* 2014;218:66-72.
- 33. Prin M, Wunsch H. The role of stepdown beds in hospital care. *Am J Respir Crit Care Med* 2014;190:1210-6.
- 34. Khan RM, Ajuaid M, Ageel H, et al. Introducing the Comprehensive Unit-Based Safety Program for mechanically ventilated patients in Saudi Arabian intensive care units. *Ann Thorac Med* 2017;12:11-6.
- Timmel J, Kent PS, Holzmueller CG, et al. Impact of the comprehensive unit-based safety program on safety culture in a surgical inpatient unit. *Jt Comm J Qual Patient Saf* 2010;36:252-60.
- Weaver SJ, Lofthus J, Sawyer M, et al. A collaborative learning network approach to improvement: the CUSP learning network. *Jt Comm J Qual Patient Saf* 2015;41:147-59.