Supervisor: Martin Gerdin Wärnberg

Co-supervisor: Jonatan Attergrim, Kelvin Szolonsky

Department of Global Public Health, Karolinska Institute and Perioperative Medicine and Intensive Care, Karolinska University Hospital Solna  
Elin Sun Cao  
Study Program in Medicine KI  
Degree project 30 credits  
Spring 2024

**Opportunities for improvement in adult trauma patients admitted to the intensive care unit: A registry-based study**

**Author:** **Elin Sun Cao**



**Förbättringsmöjligheter i vården av vuxna traumapatienter som**

**vårdats på intensivvårdsavdelningen: En register-baserad studie**

*Bakgrund:* Trauma är en ledande orsak till mortalitet och morbiditet i världen. En hörnsten i förbättringsprogram för trauma är multidisciplinära mortalitets- och morbiditetskonferenser. Syftet med dessa konferenser är att identifiera förbättringsmöjligheter (OFI). Många patienter med svårt trauma blir inlagda på intensivvårdsavdelningen (IVA). Trots detta är kännedomen om OFI hos denna patientgrupp begränsad. *Syfte*: Karaktärisera OFI hos vuxna traumapatienter som vårdats på IVA och bedöma hur patient- och processfaktorer är associerade med OFI hos denna patientgrupp. *Metod:* Detta är en registerbaserad studie som inkluderar traumapatienter mellan 2014–2023 som vårdats på Karolinska Universitetssjukhuset i Solna, varit inlagda på IVA och som har granskats avseende förekomsten av OFI vid en mortalitets- och morbiditetskonferens. Bivariabel och multivariabel logistisk regression användes för att bedöma sambandet mellan förekomsten av OFI och utvalda patient- och processfaktorer. *Resultat:* OFI identifierades hos 143 (9,9%) av 1449 patienter. Högre Revised Trauma Score (RTS) (OR 1.22; 95% CI 1.10, 1.36; p<0,001)*,* > 7 dagar på intensivvårdsavdelningen (IVA) (OR 1.83; 95% CI 1.21, 2.75; p=0.004) och överlevnad 30 dagar efter sjukhusinläggning (OR 1.85; 95% CI 1.03, 3.50; p=0.049) var signifikant associerade med högre odds för OFI i multivariabla analyser. *Slutsats:* Flera patient- och processfaktorer var associerade med ökade odds för OFI. Resultaten påvisade att det var de patienter som hade högre RTS, patienter som varit inlagda > 7 dagar på IVA och som överlevde 30 dagar efter sjukhusinläggning hade högre odds för OFI.

**Opportunities for improvement in adult trauma patients admitted to the intensive care unit: A registry-based study**

### Background: Trauma is a leading cause of mortality and morbidity worldwide. A cornerstone in trauma quality improvement programs is multidisciplinary mortality and morbidity conferences. The purpose of these conferences is to identify opportunities for improvement (OFI). Many patients with severe trauma are admitted to the intensive care unit (ICU), but little is known about OFI in this group of patients. Aim: Characterize OFI in adult trauma patients admitted to the ICU and assess how patient and process factors are associated with OFI in these patients. Methods: A registry-based study of trauma patients admitted to the ICU between 2014-2023 at the Karolinska University Hospital in Solna, Sweden, who were reviewed regarding the presence of OFI in a mortality and morbidity conference. Bivariable and multivariable logistic regression was used to determine associations between selected patient and process factors and OFI. Results: OFI was identified in 143 (9,9%) out of 1449 patients. RTS (OR 1.22; 95% CI 1.10, 1.36; p<0.001), > 7 days in the ICU (OR 1.83; 95% CI 1.21, 2.75; p=0.004) and survival 30 days after hospitalization (OR 1.85; 95% CI 1.03, 3.50; p=0.049) were significantly associated with increased odds of OFI. Conclusion: Several patient and process factors were found to be associated with OFI, indicating that patients with more days spent in the ICU, patients with higher RTS scores, and patients who were alive 30 days after hospitalization had higher odds of OFI.

# Abbreviations

ASA score = American Society of Anesthesiologists score

CT = computed tomography

GCS = Glasgow Coma Scale

ISS = Injury Severity Score

OFI = Opportunity For Improvement

RR = respiratory rate

RTS = Revised Trauma Score

SBP = systolic blood pressure

SweTrau = Swedish Trauma Registry

TQIP = Trauma Quality Improvement Programs

# Introduction

Trauma — the clinical entity of external injury and the body’s associated response — is a leading cause of mortality and morbidity worldwide. According to the Global Burden of Disease Study 2019, injury-related deaths constitute 7.6% of all deaths and take the lives of 4,3 million people yearly worldwide - more than HIV/AIDS, malaria, maternal mortality, and tuberculosis combined. The top three out of five causes of death in people aged 5-29 are injury-related: road traffic injuries, homicide, and suicide. Each year, tens of millions of people suffer non-fatal injuries leading to treatment at hospitals in emergency departments and acute care visits (4). These injuries often lead to temporary or permanent disability and the need for medical and mental rehabilitation. Disability-adjusted life-years (DALYs) is a way to measure post-discharge quality of life. Trauma accounts for one-tenth of DALYs, most of the burden in low- and middle-income countries (2). Trauma costs approximately 3% of gross domestic product yearly globally, as it mostly affects the working population (3). Trauma exposure at a young age increases the risk for mental illness and suicide leading to increased risk of an unhealthy lifestyle associated with smoking, alcohol and substance abuse, chronic disease, and cancer as well as societal problems such as crime, poverty, and violence. The burden of disease from trauma varies depending on certain conditions: socioeconomic factors, sex, age, and country. Trauma is more common in men compared women. Across all age groups worldwide, the top three leading causes of injury in males are road traffic injuries, homicide, and suicide; whereas the top three leading causes of injury in women are road traffic injuries, falls, and suicide (5).

In Sweden, the ongoing advancement of quality improvement systems places significant emphasis on addressing trauma. A cornerstone of this effort is the Swedish Trauma Registry (SweTrau), to which numerous hospitals across the country report data on trauma patients. SweTrau serves as a vital resource for continued research on trauma patients contributing to efforts to optimize trauma care nationwide. According to SweTrau, trauma is one of the leading causes of death among the working population in Sweden. Many patients survive but with severe sequelae. The leading causes of trauma in Sweden include road traffic injuries, falls, weapons, stabs, and blows. In the university hospitals of Sweden – among patients with severe trauma, 17% were deceased, 20% suffered severe disability, 49% suffered moderate disability and 14% recovered well (6).

Preventability is often defined in studies as an event that would not have occurred had the patient received ordinary standards of care, such as timely and accurate management of hypoxemia and hemorrhage during early patient management (10). The term “Opportunities for improvement” (OFI) is a term used by the World Health Organization (WHO) in their quality improvement guidelines, refers to preventable errors in care that have a direct adverse effect on patient outcomes or deviate from safe clinical practice. Quality improvement requires a feedback loop where patterns of the care of patients are analyzed consistently in an objective way. Utilizing OFIs serves as a means of identifying areas of improvement and further elevating the quality of care (7).

Albaaj et al. recently analyzed OFI in relation to different patient and process factors based on a selection of data from SweTrau including the patients admitted to the Karolinska University Hospital Solna. The study indicated that patients with moderate to severe trauma are at the highest odds of OFI (8).

For all injuries, quality hospital care can reduce short- and long-term disability. Therefore, improving planning, access, and organization of trauma care systems involving prevention, pre-hospital care, intra-hospital care, rehabilitation as well as research and quality improvement, all play a central role in reducing the overall impact after trauma (43).

## Trauma quality improvement programs

Trauma Quality Improvement Programs (TQIP) emerged in high-income countries to enhance the quality of in-hospital trauma care. Its implementation has contributed to significant improvements in the quality of care and outcomes (13). Among TQIP interventions, trauma registries along with mortality and morbidity conferences have been highlighted as key in-hospital TQIP interventions (14). In addition to these, audit filters is another intervention that is commonly a part of TQIP. All these interventions have become integral components of TQIP, facilitating ongoing improvements in trauma care (42). The purpose of multidisciplinary mortality and morbidity reviews is to discuss preventable deaths and identify OFIs in trauma care regarding structure as well as clinical processes. This way, identifying OFIs will guide us on what to focus specific efforts on. Mortality and morbidity reviews are a key method of addressing all necessary components of trauma improvement (11). Audit filters are descriptions of specific actions that should be taken, time frames in which tests or treatments should be provided, or outcomes that are expected to occur in injured patients. The purpose is to assess patients whose care falls outside the frames of these audit filters and provide feedback on how to correct errors and improve future trauma care on a systematic level (1, 12).

The paradigm of Donabedian states that implementing proven structures and processes of care is the most effective way to improve outcomes (9). Mortality and morbidity reviews as well as audit filters can be used as means of improving outcomes in line with the paradigm of Donabedian. Examples of structure and process interventions in the intensive care unit (ICU) are increased staff coverage and treatment bundles to recognize and treat common conditions such as bundles for sepsis and ventilator-associated pneumonia (10).

## Preventability among trauma patients in the intensive care unit

In the evaluation of trauma patients, the primary focus is on conducting a rapid initial assessment, known as the primary survey, to identify and address any life-threatening injuries. This is typically followed by a more comprehensive secondary survey to thoroughly assess for additional injuries. However, the secondary survey may be overlooked during the initial evaluation in the emergency department, resulting in ICU personnel assuming responsibility for conducting this assessment, along with initiating early patient management. Early management of trauma patients has over time increasingly shifted to the ICU. The goal of early management is to restore homeostasis while watching for complications, failure of procedures, and management as well as missed injuries (50). The most common challenges in trauma patient management in the ICU are airway and ventilator management, hemorrhage, transfusions, and coagulopathy (44).

Challenges of airway management in ICU-admitted trauma patients involve harmful effects of mechanical ventilation, ventilator-associated pneumonia (VAP), and severe hypoxemia due to aspiration, pneumonia, acute respiratory distress syndrome, or pulmonary contusions. Mechanical ventilation can be managed with a lung-protective strategy to reduce risks of harmful effects, there are ventilator bundles to decrease the rate of VAP, and severe hypoxemia can be treated with strategies to improve oxygenation, such as airway pressure release ventilation, pulmonary vasodilators, or extracorporeal membrane oxygenation (45, 46, 47, 48).

Hemorrhage in trauma patients involves blood loss, tissue ischemia as well as inflammatory responses. An increase in lactate and base deficit or failure to normalize them, are linked to higher mortality rates. Managing hemorrhagic shock in trauma patients typically necessitates massive transfusions (49).

Assessing preventable deaths among ICU-admitted patients poses a significant challenge due to their complex medical backgrounds, making it difficult to establish the standard of care for each individual. Moreover, despite receiving optimal treatment, these patients remain at elevated risk for complications and increased mortality rates (10).

There are many studies on OFI in relation to preventable deaths in trauma patients treated in the ICU both regarding medical and human aspects. The primary medical causes of death after trauma are hemorrhage, sepsis/multiorgan failure, and central nervous system injury. Delayed hemostatic procedures and transfusions are common areas for improvement in the hospital stage (17). The most prominent OFI regarding hemorrhage was decision-making compared to errors in technical skill. The OFIs are frequently involved in the decision between radiology, surgery, and further investigation (18).

A study from The University of Pennsylvania looked at the human aspect of OFIs in the ICU. The ICU of this hospital uses telemedicine and telemonitoring to review critical patient events. The areas of improvement they identified were team dynamics and communication, use of best practices, increased and standardized access to emergency resources, and improvement of procedural technique (19). Another study looked at how critical information was handled in the ICU and found that laboratory values and test results were the most frequently lost items. This points to the importance of communication and handling of critical information at handoffs of care (20).

Many patients with severe trauma are admitted to the ICU, but little is known about OFI in this group of patients. This is due to the complexity of this cohort making identification of OFIs in this group more challenging. While numerous studies have explored various facets of trauma care enhancement, research specifically centered on OFIs in ICU-admitted trauma patients remains limited.

## Aim

This study aims to characterize OFI in adult trauma patients admitted to the ICU and assess how patient and process factors are associated with OFI in these patients.

# Methods

## Study design

We conducted a retrospective single-center cohort study based on data from the Karolinska University trauma registry, which is part of the Swedish Trauma registry (SweTrau), and the Karolinska trauma care quality database.

## Setting

The Karolinska University Hospital in Solna is a level 1 equivalent trauma center designated to receive all severely injured patients in Stockholm (22). The hospital has direct access to radiology, surgery, emergency medicine, intensive care, and consultants from relevant specialties (23). All patients are included in a mortality and morbidity screening process, combining audit filters and an individual review by specialized nurses. Those identified as having a higher likelihood of OFI are discussed in a multidisciplinary conference held every 6-8 weeks. Examples of OFI include the need for improved organization or greater involvement of senior staff in trauma care. The presence or absence of OFI is decided among the participants of the conference and subsequently documented in the trauma care quality database.

## Participants

The trauma registry includes all patients admitted with trauma team activation, regardless of Injury Severity Score (ISS), as well as patients admitted without trauma team activation but found to have an ISS of more than 9. We included all patients who had been included in the morbidity and mortality screening process between January 1, 2014, and February 1, 2023, and who were also admitted to the ICU. We excluded patients who were younger than 15 years and patients who were dead on arrival. Patients under 15 years of age were not included in the study as they are included in other registries, and because the clinical approach for children varies from that of adults.

## Variables and data sources/measurements

### **Study outcome**

The study’s outcome was the presence of OFI — defined as “Yes” for at least one identified OFI or “No” for no identified OFI — based on consensus reached during the mortality and morbidity conference.

## Patient and process factors

The variables selected for analysis were patient and process factors chosen from the trauma registry, based on the locally used audit filters, standard epidemiological factors, and factors registered in SweTrau. Categorical factors included sex, mechanical ventilation, on-call hours, ICU length of stay, ASA preinjury, and 30-day mortality. Sex was categorized as male sex or female sex. Mechanical ventilation was stratified into “No intubation”, “Mechanical ventilation 1-7 days”, “Mechanical ventilation > 7 days” and “Unknown”. On-call hours was defined as arrival to the hospital before 8.00 a.m., after 5 p.m., or during weekends – Saturday or Sunday. ICU length of stay was categorized as “≤ 7 days” or “> 7 days”. ASA preinjury was divided into “ASA 1-2” and “ASA 3-7”. 30-day mortality was determined based on whether the patient was deceased or alive 30 days after hospitalization.

The continuous variables in the analysis were age, Revised Trauma Score (RTS), ISS, and time to first computed tomography (CT). In instances where hospital values for systolic blood pressure (SBP), respiratory rate (RR), or Glasgow Come Scale (GCS) scores were missing from the emergency department records, prehospital values were used instead. If the patient was intubated, it was assumed that their GCS was 3 and RR was 0. Subsequently, for the RTS variable, each of the variables GCS, SBP, and RR were converted RTS codes ranging from 0 to 4 based on their respective values. RTS was then calculated using the formula for RTS: 0.9368 \* GCS-code + 0.7326 SBP-code + 0.2908 RR-code to derive the RTS value.

## Statistical methods

When conducting logistic regression analysis, the number of events per variable (EPV) equal to 10 is a rule of thumb. If the EPV falls below this threshold, the results for regression analyses should be interpreted with caution (25). Keeping this guideline in mind, we selected independent variables for the analysis. To achieve this, we converted RTS, ISS, and time to first CT into continuous variables, while categorical variables were recoded into as few levels as possible without compromising the quality of the analysis.

A complete case analysis was conducted after handling missing values. We presented sample characteristics using descriptive statistics. Pearson’s Chi-squared test was used to calculate p-values for categorical variables, while the Wilcoxon rank sum test was used for continuous variables. Continuous variables were compared based on their mean values and standard deviations, while categorical variables were assessed by the number of patients within each subgroup and the corresponding subgroup percentages.

Bivariable logistic regression was used to determine unadjusted associations and multivariable logistic regression to determine adjusted associations between patient and process factors and OFI. Odds ratios (OR) with associated 95% confidence intervals were used with a 5% significance level. For continuous variables, ORs were derived by evaluating each unitary step increment. All statistical analyses were conducted using R (21).

## Ethical considerations

There are several ethical aspects to take into consideration regarding this study: voluntary participation, informed consent, anonymity, confidentiality, potential for harm, and results communication.

All patients with trauma team activation or high ISS scores are automatically included in the database used for this study. However, severe trauma may not be information patients, or their families wish to be publicly available, given that the collected data include sensitive details such as vital parameters, outcomes, type of trauma, and time of hospital arrival. Thus, the ethical principles of voluntary participation and informed consent must be considered. National databases collect data from all registered patients, often without explicit consent, which presents a drawback in terms of voluntary participation.

To respect anonymity and confidentiality, statistical analysis was initially conducted on synthetic data before being applied to actual data from the trauma registry and the trauma care quality database. This measure protects personal numbers and associated information, potentially alleviating discomfort among patients and their families.

Since this is an observational study without interventions, there is no direct potential for harm to participants. As for results communication, research findings and compiled original data are publicly accessible. While individual participants may not directly benefit, such research significantly contributes to the advancement of hospital care in Sweden, enhancing efficiency and equity. Studies conducted using data from the Karolinska University Hospital in Solna aim to improve local trauma care systems, thereby benefiting society by reducing trauma mortality rates and enhancing overall patient safety.

Based on these factors, it can be concluded that the benefits of the study outweigh the drawbacks. The study's advantages are the contribution to trauma system developments which long-term improves patient safety and equality, and the protection of the individual’s anonymity and confidentiality using synthetic data. The potential emotional harm from non-voluntary participation is acknowledged as a limitation.

The study was approved by the Swedish Ethical Review Authority, approval numbers 2023-02975-02, 2021-03531, 2021-02541.

# Results

## Participants

En bild som visar text, skärmbild, Teckensnitt, diagram

Automatiskt genererad beskrivning

A total of 14,022 patients were included in the trauma registry and trauma care quality database between 2017 and 2023, out of which 12,278 patients were excluded. Out of all patients, 11,343 were excluded because they were not treated in the ICU, three patients were excluded because they were under 15 years old, six patients were excluded because they were dead on arrival, and 928 patients were excluded because there was no data on the presence or absence OFI. This left us with a total of 1,742 patients eligible for the study. Out of these, 293 patients were excluded because there was missing data, leaving a total of 1,449 patients included in the study.

## Descriptive data

| **Table 1.** Sample characteristics. | | | | |
| --- | --- | --- | --- | --- |
| **Characteristic** | **No opportunity for improvement**, N = 1,306[[1]](#footnote-1) | **Opportunity for improvement,** N = 143 | **Overall,** N = 1449 | **p-value[[2]](#footnote-2)** |
| **Sex** |  |  |  | 0.3 |
| Female | 323 (25%) | 30 (21%) | 353 (24%) |  |
| Male | 983 (75%) | 113 (79%) | 1,096 (76%) |  |
| **Age** | 48.35 (20.86) | 50.10 (20.86) | 48.53 (20.95) | 0.3 |
| **Mechanical ventilation** |  |  |  | **0.009** |
| No intubation | 747 (57%) | 99 (69%) | 241 (17%) |  |
| Mechanical ventilation 1-7 days | 230 (18%) | 11 (7.1%) | 0.78, 2.13 |  |
| Mechanical ventilation > 7 days | 101 (7.7%) | 12 (8.4%) | 113 (7.8%) |  |
| Unknown | 228 (17%) | 21 (15%) | 249 (17% |  |
| **Revised Trauma Score** | 5.66 (1.89) | 6.29 (1.82) | 5.72 (1.90) | **<0.001** |
| **Injury Severity Score** | 23.91 (14.94) | 24.21 (11.10) | 23.94 (14.60) | 0.3 |
| **Time to first CT** | 79.66 (244.66) | 60.77 (87.38) | 77.79 (233.94) | **0.022** |
| **On call hours[[3]](#footnote-3)** | 939 (72%) | 1. 2%) | 1,042 (72%) | > 0.9 |
| **Days in the ICU** |  |  |  | **0.005** |
| > 7 days | 333 (25%) | 52 (36%) | 385 (27%) |  |
| ≤ 7 days | 973 (75%) | 91 (64%) | 1,064 (73%) |  |
| **ASA preinjury** |  |  |  | 0.3 |
| ASA 1-2 | 979 (75%) | 101 (71%) | 1,080 (75%) |  |
| ASA 3-7 | 327 (25%) | 42 (29%) | 369 (25%) |  |
| **30-day mortality** |  |  |  | **0.007** |
| Alive | 1,036 (79%) | 127 (89%) | 1,163 (80%) |  |
| Dead | 270 (21%) | 16 (11%) | 286 (20%) |  |

Table 1 present sample characteristics. Of all patients, males constituted 1,096 (76%) and females constituted 353 (24%) patients. There were 846 (58%) patients who were not intubated, 241 (17%) patients who received mechanical ventilation for 1-7 days, and 113 (7,8%) patients who received mechanical ventilation for more than 7 days. The number of patients who arrived outside working hours was 1,042 (72%). A total of 1,064 (73%) patients were equal to or less than 7 days in the ICU, and 385 (27%) patients were more than 7 days in the ICU. A total of 1,080 (75%) patients had an ASA score of 1-2 before arrival to the hospital, and 369 (25%) patients had an ASA score of 3-7 before arrival to the hospital. Amongst our patients, 30-day mortality was at 286 (20%).

## Main results

| **Table 2**. Unadjusted and adjusted logistic regression analyses of associations between patient level factors and opportunities for improvement. | | | | | | | |
| --- | --- | --- | --- | --- | --- | --- | --- |
|  |  | **Unadjusted** | | | **Adjusted** | | |
| **Characteristic** | **N** | **OR[[4]](#footnote-4)** | **95% CI** | **p-value** | **OR** | **95% CI** | **p-value** |
| **Sex** | 1,742 |  |  |  |  |  |  |
| Female |  | — | — |  | — | — |  |
| Male |  | 1.27 | 0.87, 1.91 | 0.2 | 1.21 | 0.80, 1.90 | 0.4 |
| **Age** | 1,742 | 1.00 | 1.00, 1.01 | 0.5 | 1.01 | 1.00, 1.02 | 0.2 |
| **Mechanical ventilation** | 1,742 |  |  |  |  |  |  |
| No intubation |  | — | — |  | — | — |  |
| Mechanical ventilation 1-7 days |  | 1.37 | 0.86, 2.14 | 0.2 | 1.30 | 0.78, 2.13 | 0.3 |
| Mechanical ventilation > 7 days |  | 0.83 | 0.36, 1.67 | 0.6 | 0.68 | 0.26, 1.52 | 0.4 |
| Unknown |  | 0.96 | 0.67, 1.37 | 0.8 | 1.04 | 0.69, 1.57 | 0.8 |
| **Revised Trauma Score** | 1,535 | 1.18 | 1.08, 1.30 | **<0.001** | 1.22 | 1.10, 1.36 | **<0.001** |
| **Injury Severity Score** | 1,741 | 1.00 | 0.99, 1.01 | 0.9 | 1.01 | 0.99, 1.02 | 0.3 |
| **Time to first CT** | 1,632 | 1.00 | 1.00, 1.00 | 0.3 | 1.00 | 1.00, 1.00 | 0.3 |
| **On call hours** | 1,742 | 1.05 | 0.74, 1.52 | 0.8 |  |  |  |
| **Days in the ICU** | 1,738 |  |  |  |  |  |  |
| ≤ 7 days |  | — | — |  | — | — |  |
| > 7 days |  | 1.52 | 1.08, 2.12 | **0.015** | 1.83 | 1.21, 2.75 | **0.004** |
| **ASA preinjury** | 1,735 |  |  |  |  |  |  |
| ASA 1-2 |  | — | — |  | — | — |  |
| ASA 3-7 |  | 1.20 | 0.84, 1.69 | 0.3 | 1.23 | 0.77, 1.94 | 0.4 |
| **30-day mortality** | 1,739 |  |  |  |  |  |  |
| Dead |  | — | — |  | — | — |  |
| Alive |  | 1.66 | 1.07, 2.68 | **0.030** | 1.85 | 1.03, 3.50 | **0.049** |

Table 2 shows the unadjusted and adjusted associations of the selected patient and process factors in relation to OFI. In the unadjusted analysis, the factors that were significantly associated with the presence of OFI were: Revised Trauma Score 1.18 (95% CI 1.08, 1.30; p<0.001), > 7 days in the ICU 1.52 (95% CI 1.08, 2.12; p=0.015) and survival 30 days after hospitalization 1.66 (95% CI 1.07, 2.68; p=0.030).

In the adjusted analysis the following factors demonstrated significant association with increasing odds of OFI: Revised Trauma Score 1.22 (95% CI 1.10, 1.36; p<0.001), > 7 days in the ICU 1.83 (95% CI 1.21, 2.75; p=0.004) and survival 30 days after hospitalization 1.85 (95% CI 1.03, 3.50; p=0.049).

Sex, age, mechanical ventilation, ISS, time to first CT, on-call hours, and ASA preinjury were not significantly associated with OFI in the unadjusted nor the adjusted analyses.

| **Table 3**. Unadjusted and adjusted logistic regression analyses of associations between patient level factors and opportunities for improvement in patients alive 30 days after hospitalization. | | | | | | | |
| --- | --- | --- | --- | --- | --- | --- | --- |
|  |  | **Unadjusted** | | | **Adjusted** | | |
| **Characteristic** | **N** | **OR[[5]](#footnote-5)** | **95% CI** | **p-value** | **OR** | **95% CI** | **p-value** |
| **Sex** | 1,395 |  |  |  |  |  |  |
| Female |  | — | — |  | — | — |  |
| Male |  | 1.32 | 0.87, 2.07 | 0.2 | 1.20 | 0.74, 2.00 | 0.5 |
| **Age** | 1,395 | 1.01 | 1.00, 1.02 | 0.14 | 1.01 | 0.99, 1.02 | 0.4 |
| **Mechanical ventilation** | 1,055 |  |  |  |  |  |  |
| Mechanical ventilation > 7 days |  | — | — |  | — | — |  |
| Mechanical ventilation 1-7 days |  | 0.93 | 0.40, 2.21 | 0.9 | 1.62 | 0.60, 4.39 | 0.3 |
| No intubation |  | 1.25 | 0.68, 2.55 | 0.5 | 1.51 | 0.56, 4.09 | 0.4 |
| **Revised Trauma Score** | 1,228 | 1.09 | 0.99, 1.21 | 0.094 | 1.10 | 0.91, 1.35 | 0.4 |
| **Injury Severity Score** | 1,394 | 1.01 | 1.00, 1.02 | 0.2 | 1.01 | 1.00, 1.03 | 0.2 |
| **Time to first CT** | 1,304 | 1.00 | 1.00, 1.00 | **0.037** | 1.00 | 1.00, 1.00 | 0.7 |
| **On call hours** | 1,395 | 1.05 | 0.72, 1.57 | 0.8 |  |  |  |
| **Days in the ICU** | 1,394 |  |  |  |  |  |  |
| ≤ 7 days |  | — | — |  | — | — |  |
| > 7 days |  | 1.24 | 0.85, 1.77 | 0.3 | 1.66 | 0.99, 2.74 | 0.051 |
| **ASA preinjury** | 1,392 |  |  |  |  |  |  |
| ASA 1-2 |  | — | — |  | — | — |  |
| ASA 3-7 |  | 1.18 | 0.79, 1.74 | 0.4 | 1.22 | 0.70, 2.09 | 0.5 |

Table 3 shows the unadjusted and adjusted associations of OFI and the selected patient and process factors among the patients who were alive 30 days after hospitalization. None of the patient and process factors were significantly associated with OFI in the adjusted analysis.

# Discussion

The key results from this study are that higher RTS, > 7 days length of stay in the ICU and survival 30 days after hospitalization were significantly associated with higher odds of OFI. Age, sex, mechanical ventilation, ISS, time to first CT, arrival during on-call hours, and ASA preinjury were not significantly associated with OFI in either the adjusted or the unadjusted analyses. In the subgroup analysis of patients who were alive 30 days after hospitalization, no patient or process factor was significantly associated with OFI in the adjusted analysis.

Our study found a significant association between higher RTS scores and higher odds of OFI. Patients with lower RTS scores are often in critical condition upon arrival — upon which emergency interventions are performed immediately. Conversely, patients with higher RTS scores, indicating relative stability upon arrival, may experience delayed interventions. This aligns with findings from Albaaj, where patients with relatively better vital parameter values, were significantly associated with increased odds of OFI in the unadjusted analysis, but not in the adjusted analysis (26). Patients with higher RTS scores may appear stable despite trauma, potentially leading to delayed emergency procedures.

Prolonged stay in the ICU were found to be significantly associated with higher odds of OFI in both the unadjusted and adjusted analyses. There are two things to consider regarding this variable. Firstly, prolonged ICU stay suggest patients were in a more severe condition, needing more recovery time. Additionally, longer ICU stays, coupled with initial severity, heighten the likelihood of errors, thus increasing odds of OFI. Conversely, shorter ICU stays may indicate patient demise before potential mistakes could occur, particularly in cases where mortality was anticipated rather than preventable. This finding is similar to previous research demonstrating prolonged ICU stays as a risk factor for increased mortality and morbidity in trauma patients (28). Furthermore, it also corresponds with a retrospective cohort indicating that lower initial illness severity and younger age are associated with earlier discharge, further supporting the link between higher odds of OFI and prolonged ICU stays (31).

The reason for prolonged ICU stays being associated with increased mortality has been investigated by previous studies suggesting that factors such as comorbidities, age, and illness severity upon admission do not fully elucidate the increased long-term mortality observed in this patient cohort. As the duration of ICU stay increases, the primary reason for ICU stay becomes less linked to the patient’s original condition and increasingly tied to persistent chronic illness accompanied by organ dysfunction (30). Consequently, this might be a potential explanation for the nonsignificance of predictors such as ASA preinjury and ISS in the analysis which typically reflect a patient’s initial health status.

Another factor associated with increased odds of OFI was survival 30 days after hospitalization. Previous research has identified an association between increased mortality and the severity of the injury upon admission (52), suggesting that patients at high risk of 30-day mortality are typically in poor health upon arrival. Consequently, the likelihood of OFI decreases as deaths may not be preventable. Therefore, survival to 30 days after the injury is associated with increased odds of OFI.

Our findings imply that patients were not subjected to differential treatment based on age or sex. This is corroborated by a study indicating that physicians’ willingness to admit patients to the ICU remains consistent regardless of whether the case describes a man or a woman (32). However, other research reveals nuances in the relationship between sex and patient outcomes. For instance, among severely injured patients aged 16-44 years, female sex is associated with lower in-hospital mortality rates. Female sex is also associated with a decreased likelihood of ICU admission (33). Conversely, male sex is associated with worse outcomes in postoperative ICU patients, with male patients consuming two-thirds of ICU resources (34). The higher rate of male ICU admissions and their association with adverse outcomes may be influenced by factors unrelated to the odds of OFI.

The preinjury ASA score was initially designed to evaluate a patient’s health status pre-surgery, offering insights into the risk of postoperative complications (37), and has been found to be a reliable score for predicting mortality after trauma, predicting readmission after traumatic injury, and classifying comorbidity in trauma patients. As highlighted earlier, prolonged ICU stay becomes decreasingly associated with a patient’s initial condition, supporting the lack of association between OFI and ASA score. Moreover, the ASA score paints a pre-trauma health picture contrasting with injury scores like RTS, which may offer a more precise assessment.

The lack of significance in on-call hours in both analyses suggests that patient’s arrival times did not influence their medical care, indicating that increasing medical resources outside regular working hours may not effectively reduce OFIs. This finding aligns with research on surgeons’ performance after night shifts, which found similar risks of adverse outcomes regardless of prior night work (51).

Similarly, the lack of significance of ISS in either analysis is consistent with studies suggesting that ISS has higher predictive value when combined with other parameters such as vital signs, comorbidities, and mechanism of injury. Our findings are further supported by research indicating that physiological models outperform anatomical models among ICU-admitted patients, with RTS being a physiological model and ISS being an anatomical one (39). Moreover, the New Injury Severity Score (NISS) has also been shown to be a superior predictor for ICU admission and prolonged ICU stay compared to ISS (40), possibly explaining the nonsignificance of ISS in our analysis.

## Strengths and limitations

One of the strengths of this study is its reliance on registry-based data, ensuring a high level of data quality and reliability. The use of pre-defined variables standardized their definitions, enhancing the potential for similar studies to replicate these findings. Additionally, the study’s specific inclusion criteria facilitated the formation of a well-defined cohort, further contributing to the reliability and reproducibility of the study.

However, several limitations need to be acknowledged. Firstly, the study size was limited. This was a single-center study and provided insights primarily into the context of the Karolinska Institute Hospital Solna. Consequently, the generalizability of the findings to other level 1 trauma centers and ICUs may be limited. Moreover, the small cohort size could potentially impact result accuracy.

Another limitation is the selection of patients for the morbidity and mortality review, which heavily relied on the local audit filters used at the Karolinska Institute Hospital Solna. The use of audit filters for patient selection for the mortality and morbidity review is associated with high false positive rates, ranging from 24% to 80% (41). Even though the use of audit filters is associated with high false positive rates, there is still a risk of misplacement where some patients with OFI may have been overlooked.

Additionally, the handling of the variables to the RTS score – namely, RR, SBP, and GCS – introduced potential sources of variability. The use of prehospital, emergency department, or intubation-based values for these variables may have affected data accuracy and reproducibility. However, the composite nature of the RTS score mitigates some of these concerns, as it provides a comprehensive assessment of the patient’s physiological status.

## Clinical implications

This study may hold potential for clinical implications, particularly considering trauma as a leading cause of death globally. While further research is needed to fully understand the clinical implications of our findings, our study contributes to the ongoing efforts to improve trauma care quality.

By identifying OFIs among trauma patients admitted to the ICU, our study offers valuable insights into areas where interventions can be targeted to improve patient outcomes. Specifically, our findings suggest that patients with higher RTS scores, prolonged stays in the ICU, and who were alive 30 days after hospitalization are at increased odds of OFIs. This highlights the importance of prioritizing these patients for closer monitoring and implementing targeted interventions to prevent errors in care.

Furthermore, our research provides valuable information about the patient demographics of the trauma center at Karolinska University Hospital in Solna. This understanding of patient characteristics can inform the development of tailored care plans and quality improvement initiatives aimed at optimizing trauma care delivery.

In summary, our study underscores the importance of continuous efforts to refine trauma care practices and prioritize patient safety. By addressing OFIs and tailoring interventions to high-risk patients, healthcare providers can strive to improve outcomes and enhance the quality of trauma care on a global scale.

### Health equity

In the context of trauma studies, much of the comparative research on OFI among trauma patients is registry-based and encompass all trauma admissions to a hospital regardless of age, sex, or sociodemographic factors.

Our study focused specifically on trauma patients admitted to the ICU at the Karolinska University Hospital Solna, including all patients with severe trauma in the Stockholm region. However, it’s important to note that the population of Stockholm may not be representative of Sweden as a whole. Karolinska University Hospital Solna is a specialized hospital, and access to similar high-quality care may not be uniform across other regions in the country. Therefore, the findings of our study may be more applicable to the population of Stockholm, rather than ICU settings in other parts of Sweden. This could potentially limit the generalizability of our results.

Furthermore, our study observed a predominance of males among the included patients. However, it’s essential to clarify that this imbalance is not a result of biased demographic selection. Instead, it likely reflects the higher incidence of males being admitted to the ICU and experiencing trauma, as indicated by the registry-based nature of our study.

## Generalizability

The study was conducted on 143 outcomes of OFI. The limited number of outcomes can be attributed to several inclusion criteria: the study focused on a single-center cohort, specifically targeting patients admitted to the ICU who were over 14 years old, not deceased upon arrival, and had recorded data on the presence or absence of OFI. While this approach ensured high internal validity by defining a clear and specific cohort of ICU-admitted trauma patients at Karolinska University Hospital Solna, it also led to compromised external validity due to the study’s restricted scope and sample size.

Despite the limitations in external validity, the findings of this study hold significant relevance for directing future efforts in trauma improvement within the ICU of this hospital. Additionally, by concentrating on a well-defined patient cohort, this research provides valuable insights into areas where targeted interventions may be most beneficial for enhancing trauma care outcomes.

## Future studies

Further research on OFI among ICU-admitted patients is needed to expand our understanding of this aspect of trauma care. The primary limitation of this study was its restricted sample size, highlighting the need for larger-scale investigations. Conducting multi-center studies encompassing extensive databases or merging multiple datasets could provide more comprehensive insights into OFI and contribute to enhancing TQIPs.

Building upon the findings of this study, future research should focus on specific variables such as ICU length of stay and the individual assessment of vital parameters like GCS, RR, and SBP, rather than relying solely on composite scores like RTS. Analyzing these vital signs separately may reveal which parameter holds greater predictive value for OFI.

Given the significant association between higher RTS scores and increased odds of OFI, attention should be directed towards trauma patients who appear stable upon admission but may deteriorate over time. Future research should aim to identify early indicators of worsening conditions in these patients, facilitating more targeted resource allocation and intervention strategies.

In contrast, ISS did not demonstrate a correlation with OFI in this study. Subsequent studies could explore alternative scoring systems such as NISS, RTS, and ASA scores, or combinations of physiological and anatomical scoring systems to determine the most effective predictor of OFI in trauma patients.

Overall, future research endeavors should strive to address these gaps in knowledge to optimize trauma care outcomes and contribute to the ongoing improvement of trauma quality improvement initiatives.

# Conclusion

This study aimed to characterize the association between OFI and patient and process factors among adult patients admitted to the ICU at the Karolinska Institute Hospital Solna. The findings revealed a significant association between higher RTS scores, ICU stay exceeding 7 days in, and survival 30 days after hospitalization with increased odds of OFI. These results suggest that efforts to enhance trauma care should particularly focus on patients initially presenting in relatively stable conditions and those with prolonged ICU stays.

# Contributions

This study has been conducted with continuous guidance and feedback from Martin Gerdin Wärnberg, Jonatan Attergrim, and Kelvin Szolonsky.

# Acknowledgments

I want to extend my greatest gratitude to my supervisor Martin Gerdin Wärnberg and co-supervisors Jonatan Attergrim and Kelvin Szolonsky who have provided me with guidance and support throughout this semester. Their extensive knowledge, deep insight into the subject, and availability and support have greatly contributed to the development of this project. Additionally, I want to express my gratitude to my friends and family who have been there contributing with encouragement and positive energy.

# References

1. Evans C, Howes D, Pickett W, Dagnone L. Audit filters for improving processes of care and clinical outcomes in trauma systems. Cochrane Database Syst Rev. 2009 Oct;(4):CD007590.

2. Haagsma JA, Graetz N, Bolliger I, Naghavi M, Higashi H, Mullany EC, et al. The global burden of injury: Incidence, mortality, disability-adjusted life years and time trends from the global burden of disease study 2013. Inj Prev. 2016 Feb;22(1):3–18.

3. Rossiter ND. Trauma-the forgotten pandemic? Int Orthop. 2022 Jan;46(1):3–11.

4. Global Health Data Exchange. 2019; Available from: <https://vizhub.healthdata.org/gbd-results/>

5. GBD 2017 Causes of Death Collaborators. Global, regional, and national age-sex-specific mortality for 282 causes of death in 195 countries and territories, 1980-2017: A systematic analysis for the global burden of disease study 2017. Lancet. 2018 Nov;392(10159):1736–88.

6. SweTrau. Svenska traumaregistret årsrapport 2022. 2022; Available from: <https://rcsyd.se/swetrau/wp-content/uploads/sites/10/2023/05/Årsrapport-SweTrau-2022.pdf>

7. Organization WH. Improving the quality of health services - tools and resources. 2018; Available from: <https://iris.who.int/bitstream/handle/10665/310944/9789241515085-eng.pdf?sequence=1>

8. David SD, Roy N, Solomon H, Lundborg CS, Wärnberg MG. Measuring post-discharge socioeconomic and quality of life outcomes in trauma patients: A scoping review. J Patient Rep Outcomes. 2021 Aug;5(1):68.

9. Donabedian A. Evaluating the quality of medical care. 1966. Milbank Q. 2005;83(4):691–729.

10. Dijkema LM, Dieperink W, Meurs M van, Zijlstra JG. Preventable mortality evaluation in the ICU. Crit Care. 2012 Dec;16(2):309.

11. Wilson DS, McElligott J, Fielding LP. Identification of preventable trauma deaths: Confounded inquiries? J Trauma. 1992 Jan;32(1):45–51.

12. Javali RH, Krishnamoorthy, Patil A, Srinivasarangan M, Suraj, Sriharsha. Comparison of injury severity score, new injury severity score, revised trauma score, and trauma and injury severity score for mortality prediction in elderly trauma patients. Indian J Crit Care Med. 2019 Feb;23(2):73–7.

13. Kapanadze G, Berg J, Sun Y, Gerdin Wärnberg M. Facilitators and barriers impacting in-hospital trauma quality improvement program (TQIP) implementation across country income levels: A scoping review. BMJ Open. 2023 Feb;13(2):e068219.

14. Mock C. WHO releases guidelines for trauma quality improvement programmes. Inj Prev. 2009 Oct;15(5):359.

15. Sathiyakumar V, Molina CS, Thakore RV, Obremskey WT, Sethi MK. ASA score as a predictor of 30-day perioperative readmission in patients with orthopaedic trauma injuries: An NSQIP analysis. J Orthop Trauma. 2015 Mar;29(3):e127–32.

16. Garcia AE, Bonnaig JV, Yoneda ZT, Richards JE, Ehrenfeld JM, Obremskey WT, et al. Patient variables which may predict length of stay and hospital costs in elderly patients with hip fracture. J Orthop Trauma. 2012 Nov;26(11):620–3.

17. Kwon J, Lee M, Jung K. National assessment of opportunities for improvement in preventable trauma deaths: A mixed-methods study. Healthcare (Basel). 2023 Aug;11(16).

18. O’Reilly D, Mahendran K, West A, Shirley P, Walsh M, Tai N. Opportunities for improvement in the management of patients who die from haemorrhage after trauma. Br J Surg. 2013 May;100(6):749–55.

19. Gold AK, Huffenberger A, Lane-Fall M, Pascual Lopez JL, Rock KC. Leveraging telemedicine for quality improvement video review of critical ICU events: A novel multidisciplinary form of education. Crit Care Explor. 2021 Sep;3(9):e0536.

20. Stahl K, Palileo A, Schulman CI, Wilson K, Augenstein J, Kiffin C, et al. Enhancing patient safety in the trauma/surgical intensive care unit. J Trauma. 2009 Sep;67(3):430-3; discussion 433-5.

21. Solna KU. R studio. 2020; Available from: <https://www.r-project.org>

22. America TCA of. Trauma center levels defined. 2014; Available from: <https://www.traumacenters.org/page/TraumaCenterLevels#Goals>

23. Solna KU. Traumamanual. 2020; Available from: <https://traumarummet.files.wordpress.com/2020/09/traumamanualen-2020.pdf>

24. Concato J, Peduzzi P, Holford TR, Feinstein AR. Importance of events per independent variable in proportional hazards analysis. I. Background, goals, and general strategy. J Clin Epidemiol. 1995 Dec;48(12):1495–501.

25. Peduzzi P, Concato J, Feinstein AR, Holford TR. Importance of events per independent variable in proportional hazards regression analysis. II. Accuracy and precision of regression estimates. J Clin Epidemiol. 1995 Dec;48(12):1503–10.

26. Albaaj H, Attergrim J, Strömmer L, Brattström O, Jacobsson M, Wihlke G, et al. Patient and process factors associated with opportunities for improvement in trauma care: A registry-based study. Scand J Trauma Resusc Emerg Med. 2023 Nov;31(1):87.

27. Chaudhary MA, Schoenfeld AJ, Koehlmoos TP, Cooper Z, Haider AH. Prolonged ICU stay and its association with 1-year trauma mortality: An analysis of 19,000 american patients. Am J Surg. 2019 Jul;218(1):21–6.

28. Hermans G, Van Aerde N, Meersseman P, Van Mechelen H, Debaveye Y, Wilmer A, et al. Five-year mortality and morbidity impact of prolonged versus brief ICU stay: A propensity score matched cohort study. Thorax. 2019 Nov;74(11):1037–45.

29. Viglianti EM, Kruser JM, Iwashyna T. The heterogeneity of prolonged ICU hospitalisations. Thorax. 2019 Nov;74(11):1015–7.

30. Darvall JN, Boonstra T, Norman J, Murphy D, Bailey M, Iwashyna TJ, et al. Persistent critical illness: Baseline characteristics, intensive care course, and cause of death. Crit Care Resusc. 2019 Jun;21(2):110–8.

31. Viglianti EM, Kramer R, Admon AJ, Sjoding MW, Hodgson CL, Bellomo R, et al. Late organ failures in patients with prolonged intensive care unit stays. J Crit Care. 2018 Aug;46:55–7.

32. Larsson E, Zettersten E, Jäderling G, Ohlsson A, Bell M. The influence of gender on ICU admittance. Scand J Trauma Resusc Emerg Med. 2015 Dec;23(1):108.

33. Pape M, Giannakópoulos GF, Zuidema WP, Lange-Klerk ESM de, Toor EJ, Edwards MJR, et al. Is there an association between female gender and outcome in severe trauma? A multi-center analysis in the netherlands. Scand J Trauma Resusc Emerg Med. 2019 Feb;27(1):16.

34. Reinikainen M, Niskanen M, Uusaro A, Ruokonen E. Impact of gender on treatment and outcome of ICU patients. Acta Anaesthesiol Scand. 2005 Aug;49(7):984–90.

35. Tran A, Mai T, El-Haddad J, Lampron J, Yelle J-D, Pagliarello G, et al. Preinjury ASA score as an independent predictor of readmission after major traumatic injury. Trauma Surg Acute Care Open. 2017 Nov;2(1):e000128.

36. Skaga NO, Eken T, Søvik S, Jones JM, Steen PA. Pre-injury ASA physical status classification is an independent predictor of mortality after trauma. J Trauma. 2007 Nov;63(5):972–8.

37. Ringdal KG, Skaga NO, Steen PA, Hestnes M, Laake P, Jones JM, et al. Classification of comorbidity in trauma: The reliability of pre-injury ASA physical status classification. Injury. 2013 Jan;44(1):29–35.

38. Ferguson SA, Paterson JL, Hall SJ, Jay SM, Aisbett B. On-call work: To sleep or not to sleep? It depends. Chronobiol Int. 2016 Apr;33(6):678–84.

39. Serviá L, Badia M, Montserrat N, Trujillano J. Gravedad en pacientes traumáticos ingresados en UCI. Modelos fisiológicos y anatómicos. Med Intensiva (Engl Ed). 2019 Jan;43(1):26–34.

40. Li H, Ma Y-F. New injury severity score (NISS) outperforms injury severity score (ISS) in the evaluation of severe blunt trauma patients. Chin J Traumatol. 2021 Sep;24(5):261–5.

41. Attergrim J, Szolnoky K, Strömmer L, Brattström O, Whilke G, Jacobsson M, et al. Predicting opportunities for improvement in trauma using machine learning. bioRxiv. 2023.

42. Cole E, Lecky F, West A, Smith N, Brohi K, Davenport R. The impact of a pan-regional inclusive trauma system on quality of care. Ann Surg. 2016 Jul;264(1):188–94.

43. WHO. Injuries and violence. 2019 Mar; Available from: <https://www.who.int/news-room/fact-sheets/detail/injuries-and-violence>

44. ATLS Subcommittee, American College of Surgeons’ Committee on Trauma, and International ATLS working group: Advanced Trauma Life Support Student Course Manual, 2012Ninth Edition. Chicago, IL, American College of Surgeons.

45. Fan E, Del Sorbo L, Goligher EC, et al.; American Thoracic Society, European Society of Intensive Care Medicine, and Society of Critical Care Medicine: An Official American Thoracic Society/European Society of Intensive Care Medicine/Society of Critical Care Medicine Clinical Practice Guideline: Mechanical ventilation in adult patients with acute respiratory distress syndrome. Am J Respir Crit Care Med 2017; 195:12531263

46. Pileggi C, Mascaro V, Bianco A, et al. Ventilator bundle and its effects on mortality among ICU patients: A meta-analysis. Crit Care Med 2018; 46:11671174

47. Dzierba AL, Abel EE, Buckley MS, et al. A review of inhaled nitric oxide and aerosolized epoprostenol in acute lung injury or acute respiratory distress syndrome. Pharmacotherapy 2014; 34:279290

48. Menaker J, Tesoriero RB, Tabatabai A, et al. Veno-venous extracorporeal membrane oxygenation (VV ECMO) for acute respiratory failure following injury: Outcomes in a high-volume adult trauma center with a dedicated unit for VV ECMO. World J Surg 2018; 42:23982403

49. Dezman ZD, Comer AC, Smith GS, et al. Failure to clear elevated lactate predicts 24-hour mortality in trauma patients. J Trauma Acute Care Surg 2015; 79:580585

50. Tisherman SA, Stein DM. ICU management of trauma patients. Crit Care Med. 2018 Dec;46(12):1991–7.

51. Govindarajan A, Urbach DR, Kumar M, Li Q, Murray BJ, Juurlink D, et al. Outcomes of daytime procedures performed by attending surgeons after night work. N Engl J Med. 2015 Aug;373(9):845–53.

52. <https://pubmed-ncbi-nlm-nih-gov.proxy.kib.ki.se/33712248/>

1. n (%); Mean (SD) [↑](#footnote-ref-1)
2. Pearson’s Chi-squared test; Wilcoxon rank sum test [↑](#footnote-ref-2)
3. Arrival at the hospital on Saturday or Sunday, or arrival at the hospital before 8 am or after 5 pm [↑](#footnote-ref-3)
4. OR = Odds Ratio, CI = Confidence Interval [↑](#footnote-ref-4)
5. OR = Odds Ratio, CI = Confidence Interval [↑](#footnote-ref-5)