

Predictors of mortality and median survival time of critically ill patients

Preditores de mortalidade e tempo médio de sobrevivência dos pacientes críticos
Predictores de mortalidad y tiempo promedio de supervivencia de los pacientes críticos

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Descriptores

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Abstract

Objective: To analyze the predictors of mortality and the average survival time of patients hospitalized in Intensive Care Units.

Methods: This is a prospective cohort, carried out from August 2018 to July 2019, in four adult Intensive Care Units (ICU) from the public and private network of the State of Sergipe. All adult patients were included, provided they had a minimum length of stay of 24 hours in the unit. The primary outcome was death. Secondary outcomes were dialysis, pressure injury, Acute Kidney Injury, need for invasive mechanical ventilation for more than 48 hours, infection, and length of hospital stay.

Results: Of the 432 patients, there was a predominance of death in male patients, older and coming from the emergency unit. The presence of heart failure, creatinine values >1.5 mg/dL at admission, diabetes mellitus, liver disease and smoking were also associated with the death outcome. As for the other predictors, the longest hospital stay, higher Sequential Organ Failure Assessment (SOFA), Simplified Acute Physiology (SAPS 3) and Nursing Activities Score (NAS) scores, in addition to the use of noradrenaline, stand out. The use of fentanyl was associated with increased survival time and the overall median survival time was 28 days.

Conclusion: The mortality predictors of patients admitted to the ICU in Sergipe were longer length of stay; the highest SOFA, SAPS-3 and NAS scores; creatinine >1.5 mg/dl on admission; use of vasopressor drugs and the need for dialysis.

Resumo

Objetivo: Analisar os preditores de mortalidade e o tempo médio de sobrevivência dos pacientes internados nas unidades de terapias intensivas.

Métodos: Coorte prospectiva, realizada no período de agosto de 2018 a julho de 2019, em quatro Unidades de Terapia Intensiva (UTI) de adultos, da rede pública e privada do Estado de Sergipe. Foram incluídos todos os pacientes adultos, desde que possuísem o tempo de permanência mínima de 24 horas na unidade. O desfecho primário foi o óbito. Os desfechos secundários foram: diálise, lesão por pressão, lesão renal aguda, necessidade de ventilação mecânica invasiva por mais de 48 horas, infecção e o tempo de internação.

Resultados: Dos 432 pacientes, houve predomínio de óbito em pacientes do sexo masculino, com idade mais avançada e procedentes da unidade de emergência. A presença de insuficiência cardíaca, valores de creatinina $>1,5$ mg/dL na admissão, diabetes mellitus, doença hepática e tabagismo também tiveram associação com o desfecho óbito. Quanto aos demais preditores, destacam-se o maior tempo de internação; maiores escores do *Sequential Organ Failure Assessment* (SOFA), *Simplified Acute Physiology* (SAPS 3) e *Nursing Activities Score*

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(NAS), além do uso de noradrenalina. O uso do fentanil foi associado ao aumento do tempo de sobrevida e o tempo médio de sobrevivência geral foi 28 dias.

Conclusão: Os preditores de mortalidade dos pacientes internados em UTI de Sergipe foram o maior tempo de internação; os maiores escores de SOFA, SAPS-3 e NAS; creatinina >1,5mg/dl na admissão; uso de drogas vasopressoras e a necessidade de diálise.

Resumen

Objetivo: Analizar los predictores de mortalidad y el tiempo promedio de supervivencia de los pacientes internados en unidades de cuidados intensivos.

Métodos: Cohorte prospectivo, realizado durante el período de agosto de 2018 a julio de 2019, en cuatro Unidades de Cuidados Intensivos (UCI) de adultos, de la red pública y privada del estado de Sergipe. Se incluyeron todos los pacientes adultos, con tiempo de permanencia mínima de 24 horas en la unidad. El criterio principal de valoración fue la defunción. Los criterios secundarios fueron: diálisis, úlcera por presión, lesión renal aguda, necesidad de ventilación mecánica invasiva durante más de 48 horas, infección y el tiempo de internación.

Resultados: De los 432 pacientes, hubo un predominio de defunciones en pacientes del sexo masculino, con edad más avanzada y procedentes de la unidad de emergencia. La presencia de insuficiencia cardíaca, valores de creatinina >1,5 mg/dL en la admisión, diabetes mellitus, enfermedad hepática y tabaquismo también estuvieron asociados con el desenlace de defunción. Con relación a los demás predictores, se destacaron el mayor tiempo de internación; mayores puntuaciones del *Sequential Organ Failure Assessment* (SOFA), *Simplified Acute Physiology* (SAPS 3) y *Nursing Activities Score* (NAS), además del uso de noradrenalina. El uso de fentanilo estuvo asociado con el aumento del tiempo de sobrevida y el tiempo promedio de supervivencia general fue de 28 días.

Conclusión: Los predictores de mortalidad de los pacientes internados en una UCI de Sergipe fueron: el mayor tiempo de internación; los puntajes más altos de SOFA, SAPS-3 y de NAS; creatinina >1,5mg/dl en la admisión; uso de drogas vasoactivas y la necesidad de diálisis.

Introduction

Intensive medicine has made great advances in recent decades, making the care provided in Intensive Care Units (ICU) have significant importance in patient survival. Thus, knowing the sociodemographic, clinical and epidemiological characteristics has been shown to be strategic and assisted in the identification of risks and definition of qualitative and quantitative interventions as a goal to improve the care provided.⁽¹⁾

Even with the large technological apparatus and new care modalities offered in ICUs, mortality is still high, ranging from 25 to 33%.⁽²⁻⁴⁾ In Brazil, this percentage is more expressive, between 9.6 and 58%.⁽⁵⁻⁷⁾ These data show the need for early analysis and interventions on predictors of mortality in ICUs.

Both mortality predictors and clinical characteristics of patients in intensive care already indicate that advanced age; the severity of the disease; length of stay in the ICU; hospital interventions, such as the use of mechanical ventilation and vasoactive drugs prior to ICU admission are risk factors for mortality.⁽⁸⁾ In addition, the number of associated comorbidities, higher baseline Simplified Acute Physiology (SAPS 2) and Sequential Organ Failure Assessment (SOFA) scores were also associated with worse outcomes.⁽⁹⁾ Sepsis from ICU admission and male gender

are also associated with mortality.⁽¹⁰⁾ It is worth noting that the SAPS II-adjusted risk of death increases by 5% for each additional point of this score.⁽¹⁰⁾ The assessment of the clinical profile of patients using SAPS and SOFA scores is crucial for implementing strategies to reduce mortality, through continuous assessment of these scores by the ICU care and management staff.

To assist in predicting ICU mortality, different severity and prognosis scores are available: those of organ dysfunction, such as the SOFA, which assesses the extent of the severity of organ dysfunction and the prognostic model of disease severity, which proposes to estimate the chance of death.⁽¹¹⁾ Assessment of these scores contributes to better management of ICU so that identifying predictors of ICU mortality associated with risk stratification, through the results of the scores, are crucial for the implementation of measures and for assessing quality of care.⁽¹²⁾

The relevance of this study consists in the assessment of variables that impact mortality in ICUs in Sergipe, which contributes to the implementation of strategies to reduce outcomes by the management of the respective sector, in order to cause positive impacts in the reduction of hospital mortality and in the management of quality of health services. Thus, the objective of this study was to analyze the predictors of mortality and the mean survival time of patients hospitalized in ICUs.

Methods

This is a prospective cohort carried out between August 2018 and July 2019, with 430 patients from four ICUs located in the state of Sergipe, in north-eastern Brazil, three of which are public referrals and one is private.

In order to provide anonymity, hospitals were coded as: H1, a medium-complexity hospital located in the central region of the state, is a reference in emergency care in the non-metropolitan region of Sergipe, which has 11 ICU beds. H2, located in the central-south region of Sergipe, consists of a teaching hospital linked to the *Universidade Federal de Sergipe*, with care on spontaneous demand and 22 ICU beds. H3, located in the capital, has access through regulation, with care organized by referred demand, with 05 ICU beds. H4 is a private institution, located in the capital, classified as large, with access by spontaneous demand, with 22 ICU beds. The ICUs were listed for the study due to the possibility of greater access for patients, both because they have a reference ICU in Sergipe, and because they have the main ICUs in the non-metropolitan region of the state.

Patients aged 18 years or older and with a minimum stay of 24 hours in the ICU were included. Patients without creatinine results, which made it impossible to classify Acute Kidney Injury (AKI) according to the criterion “Kidney Diseases: Improving Global Outcomes” (KDIGO) and those with missing or incomplete study variables, were excluded.

Data were collected by a trained team, in which at least two researchers, on a shift basis, made daily visits to the four ICUs, distributed throughout the state, for seven consecutive days, after the inclusion of patients in the study. The data collection instrument was developed by the authors themselves and was organized in the following domains: demographic data; clinical features; ICU admission support; and clinical outcomes.

Variables of interest were age, sex, race, origin and presence of comorbidities, according to the International Classification of Diseases (ICD-10). Primary outcomes were death and survival.

Secondary outcomes were discharge, dialysis, pressure injury (PI), AKI, need for invasive mechanical ventilation for more than 48 hours, cardiovascular and neurological complications (stroke), infection, length of hospital stay.

Mortality was defined as death from any cause during the ICU and hospital follow-up period and patients who survived until discharge or their transfer to another institution were classified as survivors.

The laboratory tests collected (urea, serum creatinine, electrolytes and liver profile) in the ICU routine, available in the medical records, were followed up to record the outcomes. After the seventh day of hospitalization, if patients were still hospitalized in the unit, researchers continued to monitor them until they left the ICU, either by discharge, death or transfer to another institution. However, the test records were no longer performed after the first seven days of hospitalization.

SAPS 3 and SOFA were calculated in the first 24 hours of ICU admission. The SAPS is a score used as a predictive index of mortality, in which SAPS 2 assesses clinical characteristics associated with the current context of patients in the ICU, while SAPS 3 makes an assessment with greater emphasis on the flow of patients until admission to the ICU, such as the in-hospital location of patients before admission to the ICU, length of hospital stay and the like.⁽¹³⁾ SOFA is the gold standard score for assessing mortality from sepsis.⁽¹³⁾ The Nursing Activities Score (NAS), the score used to assess the nursing workload, was calculated from the consultation of nursing records, medical prescription and patients' water balance in the last 24 hours. All NAS variables (basic activities, ventilatory support, cardiovascular, renal, neurological, metabolic and specific interventions) were of interest for this study.

Comorbidities were assessed using the Charlson Comorbidity Index, adjusted for age. This index assesses in-hospital mortality by measuring patients' clinical profile and comorbidities, such as the presence of diabetes mellitus, heart failure, history of acute myocardial infarction, stroke, and the like.⁽¹⁴⁾

Categorical variables were described by absolute and relative frequencies. Continuous variables were presented as mean and standard deviation, median and interquartile range. The hypothesis of independence between categorical variables was tested by the chi-square test. Differences in measures of central tendency were verified by the Mann-Whitney test. Mean survival times were estimated by Kaplan-Meier and differences between survival curves were tested by log-rank. Crude and adjusted risk ratios were estimated using Cox regression. The significance level adopted was 5% and the software used was R Core Team 2020.

The study was previously approved by the Research Ethics Committee of the *Universidade Federal de Sergipe* (UFS), with favorable opinion under number 2,830,187 and CAEE (*Certificado de Apresentação para Apreciação Ética - Certificate of Presentation for Ethical Consideration*) 92517018.0.0000.5546.

Results

During the study period, 430 patients were included. The comparative analysis between clinical and demographic characteristics of individuals shows that patients in the AKI group were older (67 [56-79] years vs. 57 [45-70] years, <0.001) and, for the most part, from of the emergency unit (p<0.001). The mortality found among the patients assessed was 32.1%. When comparing clinical and demographic characteristics, a statistically significant difference was observed for the variables age, unit of origin before admission to the ICU, diagnosis of heart failure, diabetes and liver disease, baseline creatinine > 1.5 mg/dl and a previous cerebral ischemic event, being higher among patients who died when compared with survivors. Moreover, in support of ICU admission, patients in the death group were more severe, demonstrated by greater need for vasoactive drugs, invasive procedures and worse SAPS III scores and Charlson comorbidity index (Table 1).

Table 2 shows the comparison between the assessed groups, where worse outcomes are observed

Table 1. Clinical-demographic characterization and admission support of assessed patients

Variables	Survival		p-value
	Death n(%)	Survivor n(%)	
Sex			
Male	70(50.7)	146(50)	0.888 ^Q
Female	68(49.3)	146(50)	
Age in years, med (IQR)	67(56-79)	57(45-70)	<0.001 ^W
Weight in kg, med (IQR)	61(51-70)	60.8(52.2-68.9)	0.788 ^W
Origin			
Emergency	74(54)	111(38.3)	<0.001 ^{QM}
Surgical center	17(12.4)	136(46.9)	
Medical clinic	41(29.9)	33(11.4)	
Surgical clinic	5(3.6)	10(3.4)	
Heart failure	23(17.4)	24(8.5)	0.007 ^Q
Previous acute myocardial infarction	20(14.9)	25(8.7)	0.055 ^Q
Hypertension	73(54.1)	139(48.1)	0.251 ^Q
Current smoker	16(11.9)	27(9.4)	0.417 ^Q
Ex-smoker	41(30.6)	69(23.9)	0.143 ^Q
Baseline creatinine > 1.5 mg/dl	39(29.5)	39(13.7)	<0.001 ^Q
Diabetes	49(36)	62(21.5)	0.001 ^Q
Liver disease	18(13.4)	16(5.5)	0.005 ^Q
Previous stroke	31(23.1)	25(8.7)	<0.001 ^Q
ICU admission support			
Use of dobutamine	6(4.4)	3(1)	0.024 ^Q
Use of noradrenaline	53(38.4)	37(12.7)	<0.001 ^Q
Use of fentanyl	79(56.8)	80(27.5)	<0.001 ^Q
Use of midazolam	52(37.7)	41(14.1)	<0.001 ^Q
Nasoenteral tube	82(59.4)	99(34.4)	<0.001 ^Q
Orotracheal tube	82(59.4)	87(30.1)	<0.001 ^Q
Central venous catheter	68(49.3)	111(38.4)	0.033 ^Q
Indwelling urinary catheter	102(73.9)	223(77.4)	0.424 ^Q
Nasogastric intubation	24(17.4)	34(11.8)	0.112 ^Q
Admission SAPS 3, med (IQR)	34(27-44)	21(13-31)	<0.001 ^W
Charlson. med (IQR)	4(3-6)	3(0-4)	<0.001 ^W

n - absolute frequency; % - percentage relative frequency; MED-Median; IQR- Interquartile Range; Q- Pearson's chi-square test; Pearson's QM-Chi-Square Test with Montecarlo correction; W- Mann-Whitney test; ICU- Intensive Care Unit; SAPS-Simplified Acute Physiology Score.

among those who died, including the need for dialysis (26.6% vs. 7.9%, p<0.001), development of PI (22.5% vs. 6.3%, p<0.001) and AKI (48.9% vs. 16.2%, p<0.001), acute myocardial infarction (8.0% vs. 2.1%, p=0.003), stroke (10.9% vs. 2.1%, p<0.001), use of mechanical ventilation for more than 48 hours after admission (71.7% vs. 21.2%, p<0.001) and infection (60.6% vs. 19%, p<0.001) (Table 2).

During daily follow-up of patients for seven days in the ICU, it was possible to observe that in the death group, water balance, diuresis, serum creatinine, hemoglobin and lactate levels, as well as workload (measured by NAS) and SOFA score were worse when

Table 2. Analysis of outcomes between death and survivor groups

Variables	Survival		p-value
	Death n(%)	Survivor n(%)	
Dialysis	37(26.6)	23(7.9)	<0.001 ^o
Pressure injury	31(22.5)	18(6.3)	<0.001 ^o
Stage of pressure injury			
Stage 1	8(26.7)	4(22.2)	0.053 ^{QM}
Stage 2	12(40)	8(44.4)	
Stage 3	1(3.3)	5(27.8)	
Stage 4	7(23.3)	1(5.6)	
Unclassifiable	2(6.7)	0(0)	
Acute Kidney Injury	68(48.9)	47(16.2)	<0.001 ^o
KDIGO			
KDIGO I	25(36.8)	18(38.3)	0.534 ^{QM}
KDIGO II	11(16.2)	11(23.4)	
KDIGO III	32(47.1)	18(38.3)	
Acute myocardial infarction	11(8)	6(2.1)	0.003 ^o
Stroke	15(10.9)	6(2.1)	<0.001 ^o
Mechanical ventilation ≥ 48 hours	99(71.7)	62(21.2)	<0.001 ^o
Infection	83(60.6)	55(19)	<0.001 ^o
Focus			
Pulmonary	56(69.1)	40(70.2)	0.194 ^{QM}
Urinary	7(8.6)	5(8.8)	
Blood	3(3.7)	2(3.5)	
Surgical wound	1(1.2)	5(8.8)	
Others	14(17.3)	5(8.8)	
ICUL, med (IQR)	12(6-22)	4(2-9)	<0.001 ^W
Length of stay, med (IQR)	18(9.5-33.5)	12(7-27)	0.001 ^W
Discharge/death SOFA, med (IQR)	5(4-9)	0(0-2)	<0.001 ^W
SAPS 3 of discharge/death, med (IQR)	37(27-50)	15(9-22.5)	<0.001 ^W
Discharge/death NAS, med (IQR)	59.2(50-73.5)	48.1(41-54.9)	<0.001 ^W

N - absolute frequency; %- percentage relative frequency; MED- Median; IQR- Interquartile Range; Q- Pearson's chi-square test; QM- Pearson's chi-square test with Montecarlo correction; W- Mann-Whitney test; MED- Median; IQR- Interquartile Range; ICU - Intensive Care Unit length of stay; SAPS - Simplified Acute Physiology Score; SOFA- Sequential Organ Failure Assessment; NAS- Nursing Activities Score; KDIGO- Kidney Disease: Improving Global Outcomes

compared to the survivor group. For the variables serum creatinine and urine output, in all seven days the values were significantly higher in the death group when compared to the non-survivors (Table 3).

The overall median survival time (MST) for the study patients was 28 days. The decrease in this time was related to previous smoking (MST=20.31, p=0.010), use of vasopressor (noradrenaline) (MST=23.25, p=0.018), need for dialysis (MST=21.31, p =0.039) and the development of AKI (MST= 20.65, p<0.001). However, the use of fentanyl increased the chance of survival by about 1.42 times. Moreover, the risk ratio for death increased threefold for liver disease, twice for AKI and increased by 51% with the use of noradrenalin.

Table 3. Relationship between the variables studied and the primary outcome

Variables	Survival		p-value
	Death n(%)	Survivor n(%)	
Water balance in ml/kg			
D1, med (IQR)	24.3(11.7-46.6)	20.1(8.2-39.7)	0.218
D2, med (IQR)	35.8(14.9-263.8)	27.8(5.8-361)	0.052
D3, med (IQR)	28.6(13.4-504)	16(-0.9-47.4)	<0.001
D4, med (IQR)	26.4(10.3-65.2)	15.3(0.6-36.6)	0.001
D5, med (IQR)	21.2(6.4-63.7)	15.1(1.3-40.9)	0.105
D6, med (IQR)	25.7(7.8-61.7)	15.1(1.4-33.2)	0.016
D7, med (IQR)	23.4(10.5-56.1)	15.7(3-39.9)	0.066
Diuresis in ml/kg/h			
D1, med (IQR)	0.6(0.3-1.2)	0.9(0.5-1.6)	0.004
D2, med (IQR)	0.8(0.4-1.3)	1(0.6-1.6)	0.007
D3, med (IQR)	0.9(0.5-1.3)	1.1(0.7-2.1)	0.003
D4, med (IQR)	0.8(0.3-1.2)	1.1(0.6-2)	0.002
D5, med (IQR)	0.8(0.4-1.5)	1.1(0.7-1.7)	0.028
D6, med (IQR)	0.9(0.3-1.7)	1.2(0.6-1.8)	0.011
D7, med (IQR)	0.6(0.3-1.3)	1(0.5-1.8)	0.015
Creatinine in mg/dl			
D1, med (IQR)	1.1(0.7-2.2)	0.9(0.6-1.3)	0.010
D2, med (IQR)	1.3(0.8-2)	1(0.7-1.4)	0.015
D3, med (IQR)	1.3(0.9-2.3)	0.9(0.7-1.4)	0.000
D4, med (IQR)	1.1(0.8-2)	0.8(0.6-1.4)	0.001
D5, med (IQR)	1.3(0.9-2.2)	0.8(0.5-1.2)	<0.001
D6, med (IQR)	1.2(0.8-2.4)	0.8(0.6-1.3)	<0.001
D7, med (IQR)	1.2(0.9-2.2)	0.9(0.6-1.2)	<0.001
Hemoglobin in mg/dl			
D1, med (IQR)	10.5(8.6-12.3)	11.9(9.8-13.7)	0.001
D2, med (IQR)	10.2(8.7-12)	11.3(9.3-12.5)	0.017
D3, med (IQR)	10.1(8.6-11.7)	10.7(9.2-12.2)	0.059
D4, med (IQR)	10(8.5-11.9)	10.4(8.9-12.3)	0.265
D5, med (IQR)	9(8.1-11.6)	10.5(8.8-12.1)	0.023
D6, med (IQR)	9.8(8.1-11.3)	10.1(8.7-12.1)	0.159
D7, med (IQR)	9(8-11.2)	10.1(8.6-11.8)	0.033
Lactate in mg/dl			
D1, med (IQR)	5.6(2-18.9)	14(4.5-21.4)	0.002
D2, med (IQR)	10.8(1.6-19.8)	14.5(1.9-20)	0.514
D3, med (IQR)	8.9(2.2-18)	17(8.9-20)	0.011
D4, med (IQR)	5.7(1.4-17)	14.5(7.8-20)	0.020
D5, med (IQR)	3.4(1.5-16.3)	15(5-20.3)	0.040
D6, med (IQR)	2.9(1.7-18.3)	15.5(10.3-20)	0.028
D7, med (IQR)	2(1.5-11.3)	15(10.5-20)	0.001
NAS			
D2, med (IQR)	49(43.8-54)	47.9(41.3-52)	0.090
D3, med (IQR)	49.2(43.8-55.1)	46.9(40.1-50.9)	0.001
D4, med (IQR)	50.6(44.1-55.4)	46.9(40.6-52.7)	0.002
D5, med (IQR)	50(46-55.8)	47.8(42.9-53.6)	0.009
D6, med (IQR)	50.8(45.4-57.2)	47.1(40.9-52.9)	0.001
D7, med (IQR)	51.1(47.5-56.8)	49.3(41.2-54.9)	0.045
SOFA			
D2, med (IQR)	4(3-7)	3(0-4)	<0.001
D3, med (IQR)	4(3-7)	2(0-4)	<0.001
D4, med (IQR)	4(3-7)	2.5(0-4)	<0.001
D5, med (IQR)	5(3-7)	3(1-5)	<0.001
D6, med (IQR)	5(3.5-8)	3(1-4)	<0.001
D7, med (IQR)	5(3-6)	3(1-4)	<0.001

n - absolute frequency; IQR- interquartile range; SOFA- Sequential Organ Failure Assessment; NAS - Nursing Activities Score

Discussion

In this study, it was possible to analyze the predictors of mortality in ICU patients, with emphasis on longer hospital stays; the highest SOFA, SAPS-3 and NAS scores; creatinine >1.5mg/dl on admission; the use of vasopressor drugs (noradrenaline) and the need for dialysis.

Most patients who had death as an outcome were male, with a mean age of 67 years, coming from the emergency unit. Regarding clinical characteristics, they had a history of heart failure, creatinine above 1.5 mg/dL on admission, diabetes mellitus, liver disease and stroke. The mean overall survival time was estimated at 28 days.

Advanced age represents a predictive factor for death, when associated with multiple comorbidities, especially with regard to diabetes mellitus, arterial hypertension and use of polypharmacy, that contribute to the development of AKI and, consequently, to the higher probability of death among intensive care patients.^(14,15) In addition to age, being a smoker, having liver disease and/or heart failure are predictors of ICU mortality. Therefore, it is necessary to use clinical scores capable of predicting the risk of death in this population, such as the SOFA and SAPS 3.^(16,17)

Another important finding of this study was the correlation between higher SAPS-3 and NAS scores with prolonged hospital stay. The origin of the emergence also contributed to the increase in SAPS 3. This finding is similar to other investigations, which showed higher SAPS 3 scores associated with increased length of stay, higher workloads and admission severity.^(18,19)

The NAS has been configured as an important tool for the nursing team sizing and its attributions in the face of critical patients. According to this measure, a nurse can take care of several patients or more than one nurse can take care of a patient. Therefore, studies that assess the NAS applicability have contributed directly to safety and quality of care. Recently, researchers revealed that the high nursing workload, assessed by the NAS at the time of discharge from the ICU, was associated with the risk of readmission.^(20,21) Thus, nursing professional

sizing is related to quality of hospital care, in order to reduce negative clinical outcomes.

Additionally, patients who died had a NAS score 1.23 times higher when compared to survivors. Variables such as the vasoconstrictor effect of noradrenaline, immobility in bed, high NAS score and decrease in the frequency of changes in patients' decubitus position, can affect worse outcomes, such as increased incidence of pressure injuries and AKI (AKI-KDIGO 2 and 3).⁽²²⁻²⁵⁾ It is known that excessive use of norepinephrine can increase serum creatinine levels and result in the need for dialysis. Therefore, noradrenaline has been associated with AKI in the presence of hypovolemia and shock.⁽²⁶⁾

Moreover, over time, there was an association between mortality and renal complications. In this study, it was observed that the profile of greater severity of patients, when identified through the NAS or SOFA score, associated with increased creatinine levels, increases the mortality of ICU patients due to AKI, which negatively impacts in patient care quality. Therefore, Vasconcelos et al. agree regarding the use of scales to predict mortality, in addition to associating age, male gender, increased creatinine, PI, high SOFA and NAS scores as greater chances for the development of AKI, also found in the present study.⁽²⁶⁾

Accurate measurement of patients' fluid balance is also a highly important variable for predicting clinical outcomes in the ICU, since positive fluid balance values were associated with the outcome of death in this study. The importance of constant assessment of direct creatinine, measured through the analysis of creatinine in urine, together with serum creatinine and urinary flow, assessed per hour, is highlighted, as the ideal to adequately identify patients at risk for the development of AKI.^(27,28) The promotion of management strategies for the early diagnosis of this problem is highlighted, in order to improve treatment strategies and then reduce mortality.^(29,30) The risk stratification for AKI is observed as a feasible activity for the nursing team, since both the water balance and the creatinine analysis are routine activities and represent indicators that predict adverse health outcomes.

The risk ratio for death was also associated with liver disease in this study. A retrospective analysis, with patients with liver disease in a Portuguese ICU, identified that alcohol consumption was the most frequent cause of cirrhosis and mortality after ICU admission (53.5%), with sepsis being the main cause of death in that population. In a Portuguese cohort, mortality was also associated with hepatic encephalopathy, renal replacement therapy, use of vasopressors, invasive mechanical ventilation and others.⁽²⁹⁾

In this study, death was associated with the development of PI (22.5% vs. 6.3%, $p < 0.001$). Corroborating these findings, a study, also developed in Sergipe, identified that, when the risk factors for the development of PI were assessed, patients with AKI had more than 3.5 times the chance of developing PI (95% CI, 1.08 - 11.65; $p = 0.036$). When assessing the length of ICU stay, it was observed that for each extra day of hospitalization patients is 3.5% more likely to develop a new PI.⁽²³⁾

Norepinephrine is also associated with AKI. Such a drug, when used in cases of hypovolemia and shock, can compromise renal function in order to increase mortality.⁽²⁷⁾ Therefore, its use should be performed with caution, since such a practice increases the incidence of AKI by 2.92 times, which compromises the prognosis of patients.⁽²⁷⁾

The risk ratio for death was also associated with liver disease in this cohort. A retrospective analysis, with patients with liver disease in a Portuguese ICU, identified that alcohol consumption was the most frequent cause of cirrhosis and mortality after ICU admission (53.5%), with sepsis being the main cause of death in that population. In the Portuguese cohort, mortality was also associated with hepatic encephalopathy, renal replacement therapy, use of vasopressors, invasive mechanical ventilation, and others.⁽²⁹⁾

However, the use of fentanyl increased the chance of survival by about 1.42 times. The relationship between fentanyl use depends on the context of analysis and risk exposure. However, Casault et al. associated fentanyl with higher mortality when compared to propofol and midazolam.⁽²⁹⁾

This study has some limitations. First, be conducted in a single region of the country, which has singularities in relation to material and social structures. Moreover, there was a lack of clinical and demographic data in medical records, which prevented a significant sample of participants. It is suggested to replicate the study, in a multicenter manner, in order to visualize the aforementioned variables in different realities.

It is noteworthy that this is the first study that assessed the severity scores and the nursing workload, through the NAS calculation in four important public and private intensive care units, located in a northeastern Brazilian state. The data presented may contribute to a multidisciplinary care guided by the analysis of specific clinical variables of patients, i.e., advanced age, for example, and also dynamic variables such as SOFA scores, NAS, urinary output, duration of mechanical ventilation, among others, which favors the consolidation of clinical reasoning and promotes the advancement of clinical and managerial practice strategies for managing patient care quality.

Conclusion

Mortality predictors of patients admitted to ICUs in the state of Sergipe were longer length of stay; the highest SOFA, SAPS-3 and NAS scores; creatinine > 1.5 mg/dl on admission; the use of vasopressor drugs (noradrenaline), and the need for dialysis. The use of fentanyl increased the chance of survival by about 1.42 times. The overall MST was 28 days and the risk ratio for death was increased by the presence of liver disease, AKI, and noradrenaline use. Patients who died had a longer hospital stay and higher SOFA, SAPS-3 and NAS scores. The present study contributes to the science of the factors that interfere in the mortality of critically ill patients, providing knowledge of the clinical profile of patients and proposing strategies for management and care interventions, in order to positively impact the survival of critically ill patients.

Collaborations

Oliveira JC, Vasconcelos GMT, Bispo LDG, Magro MCS, Fonseca CD, Pinheiro FGMS and Santana-Santos E contributed to project design, data analysis and interpretation, article writing, relevant critical review of intellectual content and approval of the final version to be published.

References

- Rong L, Hong L, Chen L, He J. Prevalence of and risk factors for thirst in the intensive care unit. *J Clin Nurs*. 2022;69(2):229–34.
- Albuquerque JM, Silva RF, Souza RF. Epidemiological profile and monitoring after discharge of patients hospitalized at intensive care unit. *Cogitare Enferm*. 2017; 22(3):e50609.
- Gershengorn HB, Harrison DA, Garland A, Wilcox ME, Rowan KM, Wunsch H. Association of intensive care unit patient-to-intensivist ratios with hospital mortality. *JAMA Intern Med*. 2017;177(3):388–96.
- Kiekkas P, Tzenalis A, Gklava V, Stefanopoulos N, Voyagis G, Aretha D. Delayed admission to the intensive care unit and mortality of critically ill adults: systematic review and meta-analysis. *Biomed Res Int*. 2022;2022:4083494.
- Aguiar LM, Martins GS, Valduga R, Gerez AP, Carmo EC, Cunha KD, et al. Perfil de unidades de terapia intensiva adulto no Brasil: revisão sistemática de estudos observacionais. *Rev Bras Ter Intensiva*. 2022 ;33(4):624–34.
- Guia CM, Biondi RS, Sotero S, Lima AL, Amorim FF. Perfil epidemiológico e preditores de mortalidade de uma unidade de terapia intensiva geral de hospital público do Distrito Federal. *Com Ciênc Saúde*. 2015;26(1/2):9–19.
- Paula MF, Vannuchi MT, Rossaneis MA, Haddad MC, Fernandes KB, Pissinati PS. Survival and mortality time of patients with long-stay hospitalizations in a hospital of high complexity supervivencia y factores asociados a la mortalidad en pacientes con hospitalizaciones de larga duración. *Enferm Foco*. 2021;12(4):682–7.
- Assis LG. Epidemiologia e avaliação da mortalidade em uma UTI mista de Sergipe utilizando o escore SAPS 3 [monografia]. Sergipe: Universidade Federal de Sergipe; 2019.
- Morkar DN, Dwivedi M, Patil P. Comparative Study of Sofa, Apache Ii, Saps Ii, as a Predictor of Mortality in Patients of Sepsis Admitted in Medical ICU. *J Assoc Physicians India*. 2022;70(4):11-2.
- Aguiar-Ricardo I, Mateus H, Gonçalves-Pereira J. Hidden hospital mortality in patients with sepsis discharged from the intensive care unit. *Rev Bras Ter Intensiva*. 2019;31(2):122–8.
- Keegan MT, Soares M. O que todo intensivista deve saber sobre os sistemas de pontuação de prognósticos e mortalidade ajustada ao risco. *Rev Bras Ter Intensiva*. 2016 ;28(3):264-69.
- Khwannimit B, Bhurayanontachai R, Vattanavanit V. Comparison of the accuracy of three early warning scores with SOFA score for predicting mortality in adult sepsis and septic shock patients admitted to intensive care unit. *Heart Lung*. 2019;48(3):240–4.
- Goldwasser RS, Lobo MS, de Arruda EF, Angelo SA, Lapa e Silva JR, de Salles AA, et al. Difficulties in access and estimates of public beds in intensive care units in the state of Rio de Janeiro. *Rev Saude Publica*. 2016;50:19.
- Bahlis LF, Diogo LP, Fuchsa SC. Índice de comorbidade de Charlson e outros preditores de mortalidade hospitalar em adultos com pneumonia adquirida na comunidade. *J Bras Pneumol*. 2021;47(01):1–6.
- Benichel CR, Meneguim S. Risk factors for acute renal injury in intensive clinical patients. *Acta Paul Enferm*. 2020; 33: e-APE20190064.
- Khajehali N, Khajehali Z, Tarokh MJ. The prediction of mortality influential variables in an intensive care unit: a case study. *Pers Ubiquitous Comput*. 2021 Feb 26:1-17.
- Costa E Silva PP, Codes L, Rios FF, Esteve CP, Valverde Filho MT, Lima DO, et al. Comparison of general and liver-specific prognostic scores in their ability to predict mortality in cirrhotic patients admitted to the intensive care unit. *Can J Gastroenterol Hepatol*. 2021;2021:9953106.
- Romano JL, Garcia PC, Silva DV, Moura BR, de Souza Nogueira L. Type of admission and nursing workload of critical patients: a cross-sectional study. *Nurs Crit Care*. 2019;24(6):387–91.
- Moreno RP, Metnitz PG, Almeida E, Jordan B, Bauer P, Campos RA, et al.; SAPS 3 Investigators. SAPS 3—From evaluation of the patient to evaluation of the intensive care unit. Part 2: development of a prognostic model for hospital mortality at ICU admission. *Intensive Care Med*. 2005;31(10):1345–55.
- Azevedo AV, Tonietto TA, Boniatti MM. Nursing workload on the day of discharge from the intensive care unit is associated with readmission. *Intensive Crit Care Nurs*. 2022;69:103162.
- Campanili TC, Santos VL, Strazzieri-Pulido KC, Thomaz PB, Nogueira PC. Incidência de úlceras por pressão em pacientes de Unidade de Terapia Intensiva Cardiopneumológica. *Rev Esc Enferm USP*. 2015;49(Spec No):7–14.
- Strazzieri-Pulido KC, S González CV, Nogueira PC, Padilha KG, G Santos VL. Pressure injuries in critical patients: Incidence, patient-associated factors, and nursing workload. *J Nurs Manag*. 2019;27(2):301-10.
- Coelho FU, Watanabe M, Fonseca CD, Padilha KG, Vattimo MF. Nursing Activities Score and Acute Kidney Injury. *Rev Bras Enferm*. 2017;70(3):475–80.
- Santos SJ, Oliveira JC, Almeida CP, Magalhães FB, Pinheiro FG, Vieira RC, et al. Occurrence of pressure injury in patients admitted to the intensive care unit. *REME Rev Min Enferm* 2021;25:e1367.
- Vasconcelos GM, Magro MC, Fonseca CD, Oliveira JC, Santos ES. Predictive capacity of prognostic scores for kidney injury, dialysis, and death in intensive care units. *Rev Esc Enferm USP*. 2021;55: e20210071.
- Santos DS, Silva JI, Melo IA, Marques CR, Ribeiro HL, Santos ES. Associação da lesão renal aguda com desfechos clínicos de pacientes em unidades de terapia intensiva. *Cogitare Enferm*. 2021;26:e73926.
- Law LS, Lo EA, Yeoh SF. Direct measurement of creatinine release over a short interval in intensive care settings. *J Crit Care Med*. 2021;25(7):800–2.
- Minja NW, Akrahi H, Yeates K, Kilonzo KG. Acute kidney injury and associated factors in intensive care units at a tertiary hospital in Northern Tanzania. *Can J Kidney Health Dis*. 2021 Jul 8;8:20543581211027971.
- Casault C, Soo A, Lee CH, Couillard P, Niven D, Stelfox T, Fiest K. Sedation strategy and ICU delirium: a multicentre, population-based propensity score-matched cohort study. *BMJ Open*. 2021;11(7):e045087.