

Nutritional status association with protein-calorie adequacy in critically ill patients admitted to an intensive care unit

Associação do estado nutricional com a adequação proteico-calórica em pacientes críticos internados em uma unidade de terapia intensiva

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ABSTRACT

Introduction: The association between nutritional status and protein-calorie adequacy in critically ill patients hospitalized in an intensive care unit was evaluated, using exclusive enteral nutritional therapy. **Methods:** Cross-sectional study of quantitative nature, carried out in a hospital intensive care unit in Guarapuava, PR, Brazil. Patients over 18 years old, of both sexes, on exclusive enteral nutrition therapy were included. Evaluation took place in the first 24 hours and after 72 hours of starting nutritional therapy. To assess nutritional status, body mass index, calf circumference classification, tetrapolar electrical bioimpedance and biochemical tests were used. In the assessment of protein-caloric adequacy, collections of the prescribed and administered volume were used. Descriptive statistics tests were performed using means, standard deviation, and relative and absolute frequencies in the SPSS version 25. **Results:** The sample consisted of 16 patients, with a mean age of 56.5 ± 17.3 . Most (75%) patients were men. According to BMI, 50% presented eutrophy, and, according to calf circumference, 68.8% presented muscle mass loss. There was no statistical difference in the analysis of body composition and phase angle between 24 and 72 hours. The nutritional recommendation needs of caloric adequacy of $\geq 80\%$ were achieved by two patients, and for protein adequacy, only 3 reached the milestone of $\geq 80\%$. **Conclusion:** Although the correlations do not show a direct association with clinical outcome, it is concluded that protein-caloric inadequacy leads to a worsening in patients' general condition, increasing chances of developing malnutrition, negatively influencing hospital clinical outcome.

RESUMO

Introdução: A associação do estado nutricional com a adequação proteico-calórica de pacientes críticos internados em uma unidade de terapia intensiva foi avaliada, usando terapia nutricional enteral exclusiva. **Método:** Estudo transversal de natureza quantitativa realizado em uma unidade de terapia intensiva de um hospital de Guarapuava, PR, Brasil. Foram incluídos pacientes acima de 18 anos, de ambos os sexos, em terapia de nutrição enteral exclusiva. A avaliação ocorreu nas primeiras 24 horas e após 72 horas de início da terapia nutricional. Para avaliação do estado nutricional, utilizou-se o índice de massa corporal (IMC), classificação da circunferência da panturrilha, bioimpedância elétrica tetrapolar e exames bioquímicos. Na avaliação da adequação proteico-calórica, utilizou-se coletas do volume prescrito e administrado. Foram realizados testes de estatística descritiva, com a utilização de médias, desvio padrão e frequências relativas e absolutas no programa *Statistical Package for Social Science* 25. **Resultados:** A amostra tinha 16 pacientes, com média de idade de $56,5 \pm 17,31$. Os pacientes eram majoritariamente homens (75%). Segundo o IMC, 50% dos pacientes apresentou eutrofia, e pela circunferência da panturrilha, 68,8% dos pacientes apresentaram perda de massa muscular. Não houve diferença estatística na análise da composição corporal e ângulo de fase entre as 24 e 72 horas. A adequação calórica $\geq 80\%$ foi atingida por dois pacientes, e para adequação proteica, apenas 3 alcançaram a meta de $\geq 80\%$. **Conclusão:** Apesar das correlações não demonstrarem associação direta com o desfecho clínico, conclui-se que a inadequação proteico-calórica induz a piora no estado geral dos pacientes, promovendo maiores chances de desenvolver desnutrição hospitalar negativamente influenciando o desfecho clínico.

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INTRODUCTION

Adequate nutrition is essential for maintenance and recovery of hospital patients. Malnutrition is a very prevalent multifactorial clinical condition, especially in the intensive care unit (ICU). It may be present before admission and/or be developed in the hospital, even in cases where the patient is well nourished at the admission time. Malnutrition can occur for reasons such as infection complications, length of stay, lack of appetite, hypercatabolism and inadequate energy intake^{1,2}. The consequences of malnutrition are important, as they lead to an increase in hospitalization time, hospital costs, morbidity, and mortality³.

Critical patients are hemodynamically unstable and need immediate care and adequate caloric-protein supply, due to the high neuroendocrine inflammatory response, hypermetabolism and catabolic stress. These processes induce energy-protein malnutrition, by depleting muscle and adipose tissue mass, which leads to increased length of stay and mortality rates^{2,4,5}.

Enteral nutritional therapy (ENT) is often used to support the energetic needs of these patients, reducing disease severity in those who are unable to eat orally. ENT is preferred because it is the second, more physiological route to nutrition^{6,7}. It is essential for patient recovery, maintaining the nutritional status, lowering infection complications, balancing the immune system and preventing malnutrition. However, the ENT must be constantly monitored, offering macro and micronutrients according to the needs of each patient and giving prioritizing adequate protein supply^{4,8}.

According to the European Society for Clinical Nutrition and Metabolism (ESPEN), an adequate supply of ENT must be performed early. It needs to be administered within 48 hours of hospitalization⁹, and for ENT administration to be satisfactory, a protein-caloric adequacy of 80% or more must be achieved¹⁰.

It is important to emphasize that the nutritional status assessment of critically ill patients is essential to define their protein caloric needs. However, there are no gold standard methods for this assessment. Because of this, it is feasible to use all available methods, allowing greater reliability at the time of the nutritional evaluation⁸.

Body composition analysis by bioelectrical impedance (BIA), dual-energy absorptiometry (DEXA), computed tomography (CT) and magnetic resonance imaging (MRI) are recommended to evaluate patients. However, there are more accessible alternatives, such as the physical examination, standard anthropometric measurements (muscle circumference, arm circumference and calf circumference), and handgrip strength. Biochemical methods can be used as biomarkers of malnutrition, such as C-reactive protein (CRP) and creatinine tests, used in chronic diseases related to inflammation, and to detect catabolism⁹.

Thus, the aim of this study was evaluating the association between nutritional status and protein-calorie adequacy of critically ill patients admitted to an intensive care unit, using exclusive enteral nutritional therapy.

METHODS

This is a cross-sectional, observational, quantitative study, carried out between August of 2022 and March of 2023. The study was conducted at the São Vicente de Paula Charity Hospital (HSVP), located in the city of Guarapuava, PR, Brazil. It is a reference hospital for 20 municipalities in the region. With prior authorization, this study was approved by Research Ethics Committee, COMEP-UNICENTRO (number 4,289,047).

For participation in the experiment, we had patients of both sexes, over 18 years old, with different pathologies and nutritional needs, and who signed the Free and Informed Consent Form (FICF) before the evaluation. For unconscious patients, the FICF was signed by the responsible family member. We excluded patients who underwent ENT in conjunction with oral feeding, patients unable to undergo the bioelectrical impedance test, pregnant women and those who did not sign the FICF.

Collection time was defined by the patients' stay analysis at the hospital, so that we could carry out evaluations at different times. Therefore, the assessment was initially performed within 24 hours of the ENT start and after 72 hours.

Due to difficulty in walking or decrease level of consciousness (DLC), measures were taken to apply formulas for weight and estimated height. Therefore, knee height (KH) was measured with the aid of a wooden anthropometric ruler, and we took measurements of arm circumference (AC), wrist circumference (WC) and calf circumference (CC) with a 2-meter inelastic soft ruler tape.

To stipulate the current weight (in kilograms), formulas by Ross Laboratories¹¹ were used. For women, body weight was equal to $(KH \times 1.01) + (AC \times 2.81) - 66.04$. For men, body weight was equal $(KH \times 1.19) + (AC \times 3.21) - 86.82$. To calculate height, we used the calculations by Chumlea et al. (1994)¹². For women, height was equal to $70.25 + (1.87 \times KH) - (0.06 \times \text{age})$, while the height for men was equal to $71.85 + (1.88 \times KH)$.

Body mass index (BMI) was calculated from the estimated height and weight, dividing weight (kg) by height square (m²). We classified the BMI according to the WHO guidelines¹³.

Body composition was assessed using tetrapolar electrical bioimpedance (RJL Systems model Quantum IV®, Clinton, Ohio, United States), which provided raw data for resistance, reactance, impedance and phase angle. The standardized phase angle (SPA) was calculated according to

the formula: [(observed FA – mean FA for gender and age) / standard deviation of FA for gender and age]. We classified the obtained values by referencing parameters for age and gender according to Barbosa-Silva (2005)¹⁴.

We also used a software (RJL Systems, Clinton, Ohio, United States) to obtain fat mass weight (FM), lean mass weight (LM), and % body fat and total body water¹⁵. We excluded data from 3 patients whose bioimpedance test result was not consistent with the range of resistance and reactance values. The SPA obtained through the bioimpedance test was used to compare the individuals, even in the two collection periods.

Percentage body fat for sex and age was classified according to Pollock & Wilmore (1993)¹⁶. Body fat was evaluated as excellent, average, below average, above average, poor and very poor.

To establish the protein requirement of each patient, the amount recommended for each clinical condition was used¹⁷. Basal metabolic rate (BMR) and total energy expenditure were calculated using the Harris & Benedict formula (1918)¹⁸, multiplying factors for activity, injury and thermal BMR.

Patients using ENT underwent an evolution protocol during the evaluation period, ranging from 15 ml/hour on the first day, 30 ml/hour on the second day and 45 ml/hour on the third day. These values were prescribed by the responsible nutritionist, who establishes the recommended evolution volume for each patient. To consider caloric and protein quota achieved, the adequacy equal or above 80% proposed by Ribeiro (2015)¹⁰ was used.

Biochemical tests used for analysis were from moment of patient's admission, obtained through electronic medical records.

Data were analyzed using descriptive statistics, using means, standard deviations (SD) and relative and absolute

frequencies. Descriptive values were expressed as mean ± SD. All p-values were two-sided. We established significance levels of ≤0.05. A Kolmogorov-Smirnov was performed to verify the normality distribution of numeric variables. Numerical variables comparison for related samples and parametric values was performed using the T-student test. The Wilcoxon test was used for non-parametric variables. Analyses were performed using the Statistical Package for Social Sciences (SPSS), version 25.0 (IBM Corporation, Armonk, New York, USA).

RESULTS

The sample consisted of 16 patients, with a mean age of 56.5 ± 17.3 (ages of 24 to 86). Out of all patients, 12 (75.0%) were male. Patients were hospitalized with different diagnoses, with a higher prevalence of stroke (50%), followed by cancer (12.5%), and other comorbidities (37.5%). Administration of enteral feeding to tube position was given via nasogastric route. For the outcome, most patients were discharged 8 (50.0%) and there were 7 deaths (43.8%).

Patients nutritional profile is shown in Table 1. Weight and BMI did not vary between the first and second data collection. Weight for the first and second data collection were 67.3 ± 17.6 kg and 67.5 ± 16.6 kg, respectively. BMI for the first and second data collection were 23.8 ± 5.8 kg/m² and 23.9 ± 5.5 kg/m² respectively. Mean height among patients was 168.6 ± 5.2 cm.

Most of the patients (81.3%) had absent bowel habits, some patients (12.5%) had regular bowel movement, and 1 (6.3%) had diarrhea. After the second evaluation, most of the patients with absent bowel habits started to have constipation (75%).

When classified by the CC, the majority (81.3%) of patients presented muscle mass loss. The mean of the first and second CC assessment was 32.9 ± 4.2 cm and 33.1 ± 3.7 cm, respectively.

Table 1 – Nutritional status profile of hospitalized patients receiving enteral nutrition therapy in a hospital in Guarapuava, PR, Brazil.

Variable	Classification	24 hours, n(%)	72 hours, n(%)
BMI	Underweight	3 (18.8)	3 (18.8)
	Normal weight	8 (50.0)	8 (50.0)
	Overweight/Obesity	5 (31.3)	5 (31.3)
Bowel habit	Absent	13 (81.3)	-
	Regular	2 (12.5)	3 (18.8)
	Constipation	-	12 (75.0)
	Diarrhea	1 (6.3)	1 (6.3)
CC Classification	Normal	5 (31.3)	3 (18.8)
	Muscle mass loss	11 (68.8)	13 (81.3)
Stress diagnosis	Low	2 (12.5)	-
	High	14 (87.5)	-

BMI: body mass index; CC: calf circumference.

Table 2 characterizes the patients' biochemical profile at admission. All patients had higher than recommended values for CRP exams. However, 62.5% patients presented urea content above the recommended value. The number of patients with higher than recommended values of creatinine were lower (43.8%). The average values were 12.3 ± 9.3 mg/dl for CRP, 52.6 ± 22.6 mg/dl for urea, and 1.2 ± 0.6 mg/dl for creatinine.

The body composition variables had no relation to fat body mass, skeletal muscle mass, intra and extracellular water, and SPA (Table 3). That was true for both evaluations ($p_{\text{body mass}} = 0.298$; $p_{\text{skeletal muscle}} = 0.443$; $p_{\text{intra water}} = 0.731$; $p_{\text{extra water}} = 0.311$; $p_{\text{SPA}} = 0.600$). The mean body fat was 16.8 ± 8.7 kg in the first assessment and 16.4 ± 9.1 kg in the second assessment.

In the first assessment of 13 patients with the analyzed body composition, 3 presented SPA changes from normal to reduced, while 2 presented SPA changes from reduced to normal.

Figure 1 shows the patients' caloric adequacy trajectory during ENT evolution. Only two patients reached the estimated caloric quota of $\geq 80\%$. The maximum percentage reached was 85%, by one of the patients on the third day, and by another patient who obtained their adequacy in the three days.

We observed that low percentages of protein adequacy were more frequent, and only 4 patients reached the estimated quota of $\geq 80\%$. The maximum percentage reached on the first day was 80%. On the second day, it was 73.6%, and in the third day, it was 88.6% (Figure 2).

Table 2 – Biochemical profile in patients receiving enteral nutrition therapy in a hospital in Guarapuava, PR, Brazil.

Variable	Classification	N	%
Hemoglobin	Below recommended	7	43.8
	Normal	7	43.8
	Above recommended	2	12.5
Hematocrit	Below recommended	5	37.5
	Normal	7	43.8
	Above recommended	3	18.8
Leucocytes	Below recommended	0	-
	Normal	9	56.3
	Above recommended	7	43.8
CRP	Below recommended	0	-
	Normal	0	-
	Above recommended	15	100
Urea	Below recommended	0	-
	Normal	6	37.5
	Above recommended	10	62.5
Creatinine	Below recommended	4	25.0
	Normal	5	31.3
	Above recommended	7	43.8

CRP: C-reactive protein.

Table 3 – Biochemical profile in patients receiving enteral nutrition therapy in a hospital in Guarapuava, PR, Brazil.

Evaluated Measures (mean \pm SD)	24 hours	72 hours	p-value
Fat body mass (%)	23.61 \pm 6.74	22.91 \pm 7.70	0.298
Skeletal muscle mass (kg)	23.46 \pm 7.63	21.60 \pm 8.34	0.443
Intracellular water (kg)	21.79 \pm 4.94	22.25 \pm 6.30	0.731
Extracellular water (kg)	17.70 \pm 5.07	18.90 \pm 5.82	0.311
SFA	-0.53 \pm 3.32	-1.31 \pm 4.03	0.600

%: percentage; kg: kilograms; SPA: standardized phase angle; SD: standard deviation.

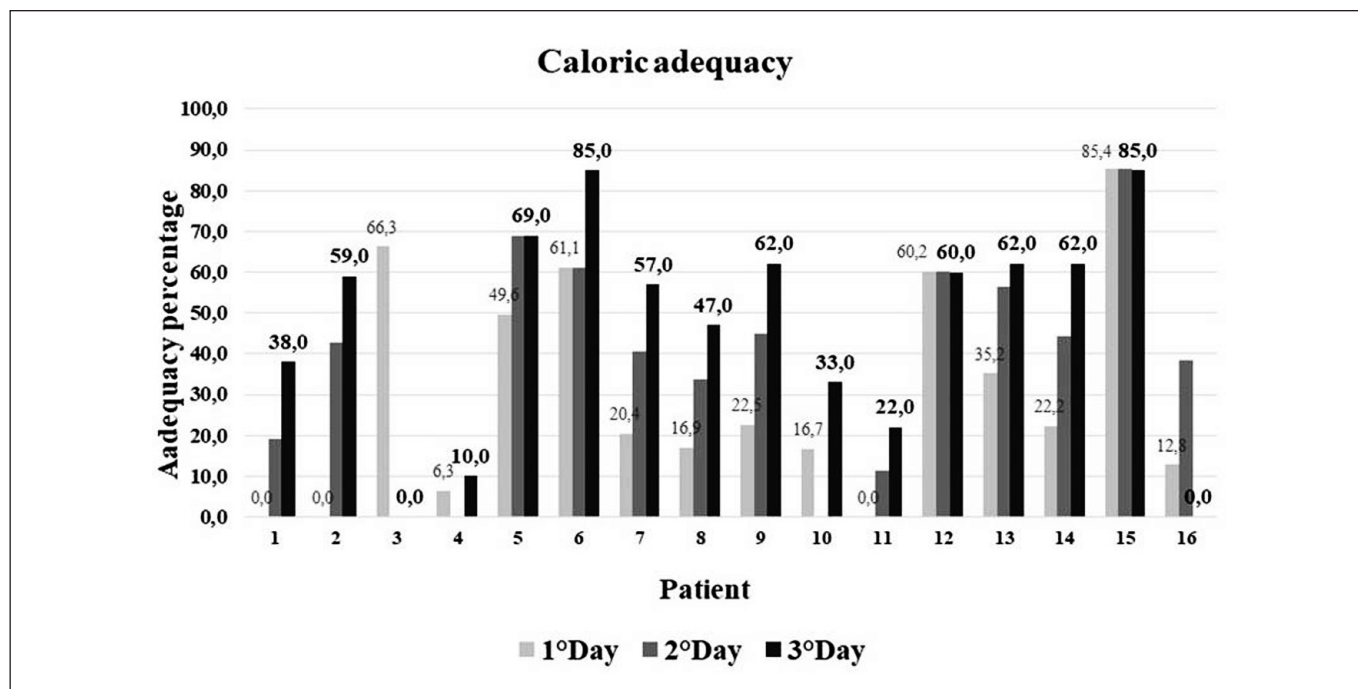


Figure 1 - Caloric adequacy of diet prescribed and ingested by patients in a hospital in Guarapuava, PR, Brazil.

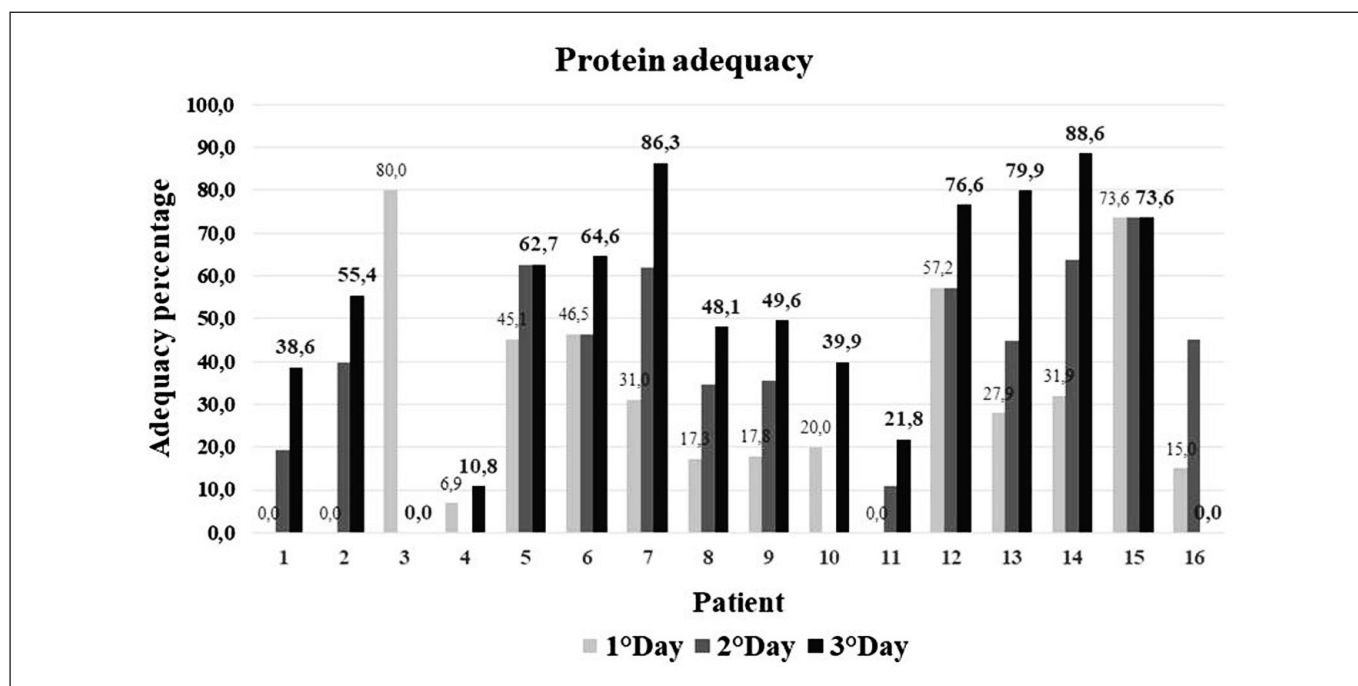


Figure 2 - Protein adequacy of diet prescribed and ingested by patients in a hospital in Guarapuava, PR, Brazil.

DISCUSSION

The different diagnoses found in this study were related to the hospital, as it is a reference center, dealing with specialties in cardiology, neurology, high-risk pregnancy, and oncology. Cerebral vascular accident (CVA) is a neurological disease responsible for most hospitalizations in hospital environments, a result also found by another study, where the hospitalization

prevalence for neurological diseases was predominant (31.0%) in relation to other comorbidities⁶.

Among male individuals, there was a greater predominance of hospitalization time, contributing to greater data collection for males. A similar result was found in another study, where most patients using ENT were male (58.7%)¹⁹. This can be explained by the low demand for health services

among men, due to feelings of diagnosis fear, shame due to the false idea of not being prone to acquiring diseases, impatience in relation to health services organization, and delay in care. These factors are also influenced by sociodemographic characteristics and lifestyle, including alcoholism, residence location, and education level²⁰.

Critical patients often have gastrointestinal problems and constipation. This is commonly caused by medication such as vasoactive drugs, enteral diets with low fiber supply, and/or electrolyte imbalance. These problems should be avoided, as the prevention of intestinal constipation avoids local irritation and lowers perforation risk. These conditions could lead to longer hospital stays, increased mortality, and intolerance to enteral nutrition^{3,21}.

There was a prevalence of constipation in the second data collection. Similarly, a study described the high intestinal complications preponderance in patients admitted to the ICU (81.1%)²¹. This can be explained due to the intestinal microbiota modulation by ENT. The treatment for intestinal imbalances is based on the use of soluble fiber, for critically ill patients who have diarrhea, and insoluble fiber, for those with constipation. It is up to the responsible team to decide the best approach for each complication²².

According to data available in a survey with 4000 hospitalized patients, 48% of these had malnutrition²³. Critically ill patients are prone to develop muscle mass depletion due to intense inflammatory and metabolic response to stress. In another study, 87.5% of patients presented high stress²⁴, which could explain the 68.8% who presented muscle mass loss due to CC in the second evaluation. However, according to BMI, most patients were eutrophic. A similar result was found in another study, that evaluated the protein adequacy of 188 critically ill patients. Here, 54.8% of the sample was classified as eutrophic⁴.

In this study, entire sample showed CRP values above the recommended, in line with the results found in another survey²⁵. However, in a study that evaluated 130 patients for 7 days, a negative relationship between CRP and infused volume was observed¹⁰. This could suggest that the metabolic alterations of the systemic evaluation would lead patients to tolerate smaller volumes of enteral nutrition, contributing for caloric inadequacy, a result that was not observed in this study. High CRP values tend to indicate neurological atrophy, which, in the long-term, influences clinical outcome, especially in those diagnosed with stroke²⁶, which many of the patients in our study had.

Creatinine is a renal function biomarker, derived from the creatine metabolism. Due to its origin in myocytes, it can be used to estimate muscle mass. In patients with critical illnesses, creatinine can increase or decrease, depending on the underlying disease. However, it can also be influenced by

medications used in the ICU. Therefore, it should be observed as a complementary measure for assessing nutritional status, and not on an individual basis²⁷. One study associated creatinine values below 0.6 mg/dl to a lower survival rate among hospitalized patients²⁷. In our study, 25% of the sample presented creatinine levels below the recommended level (<0.6 mg/dl), and the majority of patients presented high values (43.8%). This indicates muscle catabolism. A similar study evaluated the nutritional status in 31 patients in an intensive care unit, where patients also had high creatinine values⁸.

Our study also found high values of urea. This was similar to a result also found in one study that presented a high mean value of urea (72.9 ± 45.3 mg/dl)⁸, and another, which found that 19 out of 20 patients had values higher than recommended for serum urea (95%). Urea is used as an indicative measure of renal failure, and when below the recommended level, it tends to illustrate a lack of proteins in the body²⁹, but this measure can be induced by other factors not related to renal function⁸.

SPA is used as an indicative for people at high risk. Reduced SPA in critically ill patients is linked to an unfavorable prognosis¹⁴. It was observed that some patients (23.0%) started to present normal to reduced SPA in the second evaluation, which could indicate a worsening in these patients' prognosis.

Skeletal muscle mass, along with fat body mass, intra and extracellular water did not show significant changes after the second evaluation. This result shows that the collection time may be small for there to be noticeable differences in the enteral diet evolution. We found no papers that evaluated body composition of critically ill patients using ENT at different times, which limits the discussion on this subject.

In this study, the volume received from the enteral diet was evaluated for three days, and protein-caloric adequacy was verified during this time. Only two achieved caloric adequacies of $\geq 80\%$ during the evaluation period. This should be monitored, especially in critically ill patients admitted to the ICU, as studies point that better caloric adequacies lead to better adherence to ENT, lower mortality rates, and shorter hospitalization periods^{4,8}. It also prevents declines in the immune response, changes in body composition, and malnutrition. Another study observed that early ENT in critically ill patients increased the chance of hospital discharge by 1.22 times³⁰. The same benefit was also reported by another study, because ENT minimizes malnutrition, consequently avoiding complications during hospitalization⁹. It is useful to characterize early ENT as the one administered within the first 24 to 48 hours of hospitalization. In the unit where our research was carried out, there are no protocols that require early ENT. However, authors suggest that it is a necessary practice, especially in patients with malnutrition, as this helps avoiding further

reductions in nutritional status^{30,31}. Despite that, a recent meta-analysis did not indicate a decrease in hospitalization time in the ICU, time under mechanical ventilation or mortality rate among patients who had a caloric adequacy of $\geq 70\%$ ³². Considering that most patients did not receive early ENT (56.25%), clinical outcome may not be associated with the initial time of diet administration.

Furthermore, in relation to the caloric adequacy assessment, similar studies obtained positive results for adequacy. Achieving an adequacy of $>80\%$ was associated to shorter hospital stays^{6,7}. However, inadequacies found in this study corroborate a finding where, out of a total 1,122 patients, 646 presented caloric inadequacy³³.

In the study carried out with critically ill patients, the early use of ENT indicated an increase in dietary tolerance, with a prescribed volume exceeding 70%³¹. However, we did not find the same result along the diet evolution process, indicating less adherence to ENT in the long term, which could lead to a worsening in patient's general condition³¹.

Regarding protein inadequacy, considering the first day, only 1 patient reached the 80% recommendation. In the second day, no patient reached the recommendation value, reaching the maximum percentage of 73.6%. Finally, on the third day, 4 patients had adequacy $\geq 80\%$. Therefore, out of the 16 patients who received ENT, only 3 (18.7%) reached the recommendation within 72 hours. This result is similar to one study where 56.4% of the patients had insufficient adjustments⁴. Multidisciplinary teams should be more concerned with protein intake than with caloric intake, as receiving 1.3 g/kg/day of protein can improve immune response among patients⁹.

Low protein intake tends to negatively influence survival rate, increasing mortality and increasing chances of malnutrition. There is a recommendation for patients to reach recommended protein intake values within 72 hours³⁴. However, this did not occur in this study. If necessary, enteral formulas should be supplemented to meet satisfactory proteins quantities, so that each patient can reach their estimated protein quota⁸. Although clinical outcome is not associated with protein-calorie adequacy, studies suggest that this inadequacy leads to a higher mortality rate and interferes with the chances of hospital discharge^{2,8,34}.

It is also known that factors may be associated with protein-calorie inadequacy, as hospitalized patients may need to fast for surgical procedures and examinations. They might also suffer from tube complications and gastrointestinal symptoms². This highlights the importance of nutritionist participation in monitoring patients' diets and their position among the nursing team, so that they can help develop specific nutritional plans for critically ill patients. This will ensure that the proposed goal is achieved, avoiding periods of prolonged fasting and also allowing for individual meal plans.

This study has limitations, such as the time required to carry out the research and low turnover of patients using exclusive ENT, impacting the sample number.

CONCLUSION

Although correlative analyzes do not demonstrate a negative effect on body composition, it is concluded that protein-calorie inadequacy leads to worsening conditions in patients, promoting greater chances of developing hospital malnutrition, and, consequently, negatively influencing the clinical outcome. It was observed that nutritional assessment is essential for carrying out an accurate diagnosis of the nutritional status in critically ill patients and defining the correct caloric-protein supply, with the aim of avoiding or recovering malnourished patients. It is important to highlight the need to develop individual and adapted protocols for hospital service available, minimizing differences between the plans for caloric-protein supply and its implementation among critical patients, given the essentiality of nutritional support, and, if the nutrition support is not enough, its related complications.

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