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# Intravenous overload of fluids and sodium may contribute to the lower infusion of enteral nutrition in critically ill patients 

## A sobrecarga intravenosa de fluidos e sódio pode contribuir para a menor infusão de nutrição enteral em pacientes críticos

## ABSTRACT

Objective: To evaluate the effects of intravenous infusion of fluids and sodium on the first day of admission on infusion of enteral nutrition in the first 5 days in intensive care patients.

Methods: A prospective cohort study was conducted with critical nonsurgical patients admitted for at least 5 days who were on mechanical ventilation and receiving enteral nutrition. The amount of intravenous fluids and sodium infused on the first day and the volume of enteral nutrition infused in the first 5 days were investigated. The volume of intravenous fluids $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of body weight and sodium (above or below the 25th percentile) infused on the first day was compared with infused enteral nutrition.

Results: A total of 86 patients were studied, with a mean ( $\pm$ standard deviation) of $65 \pm 17$ years, of which $54.7 \%$ were female. On the first day, $3,393.7 \pm 1,417.0 \mathrm{~mL}$ of fluid ( $48.2 \pm$
$23.0 \mathrm{~mL} / \mathrm{kg}$ ) and $12.2 \pm 5.1 \mathrm{~g}$ of sodium were administered. Fifty-eight (67.4\%) patients received more than $35 \mathrm{~mL} / \mathrm{kg}$ of fluids. In 5 days, $67 \pm 19.8 \% ~(2,993.8$ $\pm 1,324.4 \mathrm{~mL}$ ) of the prescribed enteral nutrition was received. Patients who received $>35 \mathrm{~mL} / \mathrm{kg}$ of intravenous fluids also received less enteral nutrition in 5 days ( $2,781.4 \pm 1,337.9$ versus $3,433.6$ $\pm 1,202.2 \mathrm{~mL} ; \mathrm{p}=0.03$ ) versus those who received $\leq 35 \mathrm{~mL} / \mathrm{kg}$. Patients with intravenous sodium infusion above the 25 th percentile ( $\geq 8.73 \mathrm{~g}$ ) on the first day received less enteral nutrition volume in 5 days ( $2,827.2 \pm 1,398.0$ versus $3,509.3$ $\pm 911.9 \mathrm{~mL} ; \mathrm{p}=0.02$ ).

Conclusion: The results of this study support the assumption that the administration of intravenous fluids > $35 \mathrm{~mL} / \mathrm{kg}$ and sodium $\geq 8.73 \mathrm{~g}$ on the first day of hospitalization may contribute to the lower infusion of enteral nutrition in critically ill patients.

Keywords: Critical care; Fluids; Sodium; Enteral nutrition

## INTRODUCTION

Resuscitation with fluid infusion is one of the most frequent interventions performed in patients in intensive care, especially in the presence of shock. ${ }^{(1)}$ The physiological aim of resuscitation is to restore or maintain the effective circulating volume to ensure adequate tissue perfusion. ${ }^{(2-4)}$ However, excess fluid can result in adverse effects. ${ }^{(5,6)}$ Rapid redistribution of the infused volume leads to capillary damage ${ }^{(7)}$ and an increase in capillary permeability; thus, only 5 to $20 \%$ of infusions remain intravascular 90 minutes after infusion. ${ }^{(8)}$ This excess also results in sodium and water retention, with consequent acute kidney injury and impaired oxygen supply to tissues due to the low effective circulating volume. ${ }^{(9)}$ In this context, several studies have shown increased mortality with
intravenous fluid overload. ${ }^{(10-13)}$ Thus, more restrictive fluid administration strategies may be beneficial to avoid overload and increased morbidity and mortality, ${ }^{(10,14,15)}$ especially on the first day, when the patient receives more volume. It is important to note that the terms "restrictive" and "liberal" in fluid administration are not uniform. ${ }^{(10)}$ On the other hand, there is a close relationship between nutritional therapy and fluid supply. In clinical practice, nutrients, water, and electrolyte balance are interconnected during treatment, ${ }^{(11)}$ and calories and nutrient supply are crucial for reducing complications and death. ${ }^{(17,18)}$ However, in clinical practice, only 50 to $87 \%$ of the prescribed enteral diet is effectively infused. ${ }^{(19,20)}$ In this relationship between nutrients and fluids, a high infusion of crystalloid solutions results in anasarca, inadequate weight gain, ${ }^{(21)}$ intestinal loop edema, gastroparesis, vomiting, and adynamic ileus. These adverse effects, caused by an excess volume of fluids and sodium, may contribute to lower administration of an enteral diet and an increase in caloric and protein deficits. ${ }^{(18)}$ Thus, the guidelines of the Surviving Sepsis Campaign ${ }^{(22)}$ recommend that patients with hypoperfusion, hypotension or hypovolemia receive an initial volume of fluids of only $30 \mathrm{~mL} / \mathrm{kg}$. Corroborating this recommendation, the consensus Guidelines on Intravenous Fluid Therapy for Adult Surgical Patients (GIFTASUP) also recommend a restrictive intravenous fluid volume. ${ }^{(23)}$

This study aimed to evaluate the effects of intravenous administration of fluids and sodium on the first day of hospitalization with the infusion of an enteral diet in nonsurgical intensive care patients undergoing mechanical ventilation.

## METHODS

A prospective cohort study was conducted between October 2014 and December 2015 in the intensive care unit (ICU) of a private hospital (Hospital Santa Rosa) in Cuiabá (MT). The study was approved by the Ethics Committee on Research with Humans (CAAE: 37465414.0.0000.5541) and was conducted in accordance with the Declaration of Helsinki (2000). Relatives or guardians of patients signed an informed consent form (ICF).

Included patients were admitted to the ICU for at least 5 days for clinical, nonsurgical, mechanical ventilation during the first 24 hours of hospitalization and received exclusive enteral nutritional therapy (ENT). Surgical patients undergoing spontaneous breathing, pregnant women, those with late onset of nutrition (> 48 hours of
hospitalization), those who received exclusive parenteral or enteral-associated nutrition, those with hemodynamic instability, and those who died within the first 5 days of hospitalization were excluded.

The outcome variables investigated were the total volume of intravenous fluids administered on the first day of hospitalization ( mL ) in $\mathrm{mL} / \mathrm{kg}$ of body weight, grams of sodium administered on the first day of hospitalization, total volume of ENT prescribed and infused in 5 days, volume of ENT prescribed and infused on the first day, the percentage of ENT infused in 5 days (enteral feeding volume infused in 5 consecutive days $\times 100 /$ total volume of ENT prescribed in 5 days) and the protein-calorie deficit over 5 days. Considering the recommendations regarding fluid resuscitation in the acute phase ${ }^{(22,23)}$ and according to the ASPEN Board of Directors and the Clinical Guidelines Task, ${ }^{(24)}$ which recommends 30 to $40 \mathrm{~mL} / \mathrm{kg}$ of body weight, the volume of intravenous fluids was categorized as $>35 \mathrm{~mL} / \mathrm{kg} /$ day and $\leq 35 \mathrm{~mL} / \mathrm{kg} /$ day. For statistical analysis, the sodium value administered was categorized below or above the 25 th percentile $(8.73 \mathrm{~g}$ ) because this was the lowest interquartile range. The volume of intravenous fluids infused on the first day $/ \mathrm{kg}$ of body weight ( $\leq 35 \mathrm{~mL} / \mathrm{kg}$ or $>35 \mathrm{~mL} / \mathrm{kg}$ ) and the amount of sodium above or below the 25 th percentile were correlated with the total enteral diet prescribed and infused in 5 days $(\mathrm{mL})$, the volume of ENT prescribed and infused on the first day, the percentage of prescribed enteral diet infused, and the protein-calorie deficiency over 5 days.

To characterize the sample, age, patients $\geq 60$ years, sex, estimated body weight, main causes that led to hospitalization, most frequent digestive tract disorders (constipation, diarrhea, abdominal distention, and melena), volume drained by nasogastric tube (NG tube; for patients who required this procedure), Simplified Acute Physiology Score III (SAPS III) score, amount of noradrenaline ( $\mathrm{mcg} / \mathrm{kg} / \mathrm{minute}$ ), biochemical measurements (mean of 5 days of collection) of C-reactive protein (CRP; mg/L), serum albumin ( $\mathrm{g} / \mathrm{dL}$ ), lactate ( $\mathrm{mmol} / \mathrm{L}$ ) and serum glucose $(\mathrm{mg} / \mathrm{dL})$ were recorded, and the CRP/albumin ratio was calculated. The nutritional status of patients in the first 24 hours of admission and their calorie and protein requirements were also evaluated. The length of stay in the ICU and ICU mortality at 28 days were also recorded. Assessment of nutritional status was performed by the subjective global assessment (SGA A corresponded to well-nourished; SGA B, corresponded to risk of malnutrition or moderate malnutrition, and SGA C, corresponded to severe malnutrition).

## Protocol of nutritional therapy

Enteral diet was started in the first 24 hours but only in the presence of hemodynamic stability and after confirmation of the location of the probe by X-ray. Calorie and protein requirements were assessed according to the European Society for Parenteral and Enteral Nutrition (ESPEN). ${ }^{(17)} \mathrm{A}$ total of 25 to $30 \mathrm{kcal} / \mathrm{kg}$ and 1.25 to 2.0 g of protein $/ \mathrm{kg}$ of body weight were calculated. It was planned to reach the calculated need on the third or fourth day of diet. To achieve this goal, a third and a quarter of the need/day of enteral nutrition were prescribed.

## Volume of intravenous fluids and sodium

During the first 5 days of hospitalization, the volume of intravenous fluids and the amount of sodium administered were recorded. Crystalloids ( $0.9 \%$ saline solution; simple Ringer's solution, lactated Ringer's solution or glycoprotein solution), colloids, distilled water, dilution serum, drug volumes, and, lastly, blood or derivatives were considered for this purpose. The amount of sodium (grams) was determined according to the amount of crystalloid fluid administered on the first day, according to the composition of the saline solution. The researcher had no influence on the choice or method of fluid resuscitation, which was performed as the intensive care physician judged necessary.

## Statistical analysis

The chi-square test was used for the categorical variables. Continuous variables were analyzed by Levene's test to ascertain homogeneity, followed by the KolmogorovSmirnov test to determine normality. Student's t -test was used for homogeneous data with a normal distribution. Nonhomogeneous data were analyzed using the nonparametric Mann-Whitney test. The volume of infused enteral therapy ( $<$ or $\geq 2,063 \mathrm{~mL}$ ) was categorized over 5 days by the 25 th percentile. Continuous data are expressed as the mean $\pm$ standard deviation or median and variation. A significance level of $5 \%(p<0.05)$ was established. Statistical Package for the Social Sciences (SPSS) version 20.0 was used.

## RESULTS

Of 124 eligible patients, 38 were excluded because of the need for another therapy, in addition to the enteral or fasting period, and because of hemodynamic instability, examinations or procedures in the first 5 days (20),
surgical procedures (5), death (7) within the first 5 days, and family members not agreeing (6) to sign the ICF (Figure 1).

A total of 86 critical adult patients were prospectively studied, 43 ( $50 \%$ ) of whom were hospitalized due to cardiorespiratory disease, 15 ( $17.5 \%$ ) due to neurological conditions, 9 ( $10.5 \%$ ) due to neoplasms, 8 ( $9.3 \%$ ) due to trauma, such as from falling from the patient's own height, and 11 ( $12.7 \%$ ) due to other causes.

The baseline characteristics of all patients and those who received $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of fluids are provided in table 1 . Patients who received more than $35 \mathrm{~mL} / \mathrm{kg}$ of fluids had a lower estimated body weight ( $69.8 \pm 15.3$ versus $81.0 \pm 5.3 \mathrm{~kg} ; \mathrm{p}=0.002$ ), required more protein $/ \mathrm{kg}$ ( $1.39 \pm 0.16$ versus $1.30 \pm 0.18 ; \mathrm{p}=0.035$ ) and showed a greater decrease in serum albumin levels $(2.80 \pm 0.50$ versus $3.13 \pm 0.60 \mathrm{~g} / \mathrm{dL} ; \mathrm{p}=0.012$ ) compared to those receiving $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of fluids. There was no differences for the other data ( $\mathrm{p}>0.05$ ).

## Digestive tract disorders

ENT was started within 24 hours for all patients. The disorders of the digestive tract and volume drained by NG tube distributed to all patients and to those who received intravenous fluids $>$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ on the first day are provided in table 2. There was no difference between the groups ( $\mathrm{p}>0.05$ ).

## Infusion of intravenous fluids and sodium on the first day

On the first day of hospitalization, 58 (67.4\%) patients received more than $35 \mathrm{~mL} / \mathrm{kg}$ of fluids, and 28 ( $32.6 \%$ ) received $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of body weight. The amount of intravenous fluids administered on the first day was $3,393.7 \pm 1,417.0 \mathrm{~mL}$, corresponding to $48.2 \pm 23.0 \mathrm{~mL} /$ kg , with a median of $44.4(16.1-168.7) \mathrm{mL} / \mathrm{kg}$. Sixty-five ( $75.6 \%$ ) patients received a quantity of sodium $\geq 8.73 \mathrm{~g}$ on the first day (cutoff in the 25 th percentile). The mean infusion of sodium on the first day was $12.21 \pm 5.1 \mathrm{~g}$, with a median of $10.8(4.3-33.4) \mathrm{g}$.

## Intravenous fluids

Patients who received $>35 \mathrm{~mL} / \mathrm{kg}$ intravenous fluids on the first day received a lower volume of enteral diet over 5 days ( $2.781 \pm 1.338 \mathrm{~mL}$ versus $3.433 \pm 1.202 \mathrm{~mL}$; $\mathrm{p}=0.032$ ) and a lower infused percentage ( $64.3 \pm 19.7 \%$ versus $74.9 \pm 18.6 \% ; \mathrm{p}=0.020$ ) than did those who received $\leq 35 \mathrm{~mL} / \mathrm{kg}$. There were no differences for the


Figure 1 - Total number of patients eligible and excluded from the study. ICF - informed consent form.

Table 1 - Baseline characteristics of all the studied patients and, among them, those who received $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of body weight of fluids on the first day

| Variables |  | Infusion volume $>\mathbf{3 5 m L} / \mathrm{kg}$ |  | *p value |
| :---: | :---: | :---: | :---: | :---: |
|  | ( $\mathrm{n}=86$ ) | Yes ( $\mathrm{n}=58$ ) | No ( $\mathrm{n}=28$ ) |  |
| Age (years) | $64.8 \pm 17.4$ | $64.4 \pm 18.6$ | $66.3 \pm 13.1$ | 0.582 |
| Body weight (kg) | $73.4 \pm 16.2$ | $69.8 \pm 15.3$ | $81.0 \pm 5.3$ | 0.002 |
| SAPS III | $64.6 \pm 16.2$ | $67.1 \pm 17.5$ | $57.7 \pm 10.1$ | 0.165 |
| Calorie needs (kcal) | $24.14 \pm 4.0$ | $24.6 \pm 4.1$ | $23.2 \pm 3.4$ | 0.126 |
| Protein requirements (g) | $1.36 \pm 0.17$ | $1.39 \pm 0.16$ | $1.30 \pm 0.18$ | 0.035 |
| Serum albumin (g/dL) | $2.90 \pm 0.56$ | $2.80 \pm 0.50$ | $3.13 \pm 0.60$ | 0.012 |
| CRP (mg/dL) | $125.61 \pm 83.6$ | $125.7 \pm 81.9$ | $125 \pm 88.6$ | 0.982 |
| CRP/albumin ratio | $46.5 \pm 34.0$ | $48.4 \pm 34.0$ | $42.5 \pm 34.0$ | 0.464 |
| Lactate (mmol/L) | $23.7 \pm 11.1$ | $23.3 \pm 11.1$ | $24.7 \pm 11.4$ | 0.579 |
| Blood sugar (mg/d) | $181.7 \pm 50.5$ | $176.0 \pm 46.6$ | $194.0 \pm 57.2$ | 0.133 |
| Noradrenaline (mcg/kg/minute) | $0.681 \pm 0.678$ | $0.729 \pm 0.729$ | $0.582 \pm 0.555$ | 0.350 |
| Length of hospitalization (days) | $32.2 \pm 34.3$ | $34.0 \pm 39.01$ | $28.4 \pm 21.7$ | 0.496 |
| Elderly | 63 (73.3) | 42 (72.4) | 21 (75) | 0.800 |
| Female sex | 47 (54.7) | 34 (58.6) | 13 (46.4) | 0.287 |
| Nutritional status |  |  |  |  |
| SGA $=\mathrm{A}^{\dagger}$ | 9 (10.5) | 5 (8.6) | 4 (14.3) | 0.421 |
| SGA $=B^{\ddagger}$ | 64 (74.5) | 42 (72.4) | 22 (78.6) | 0.540 |
| SGA $=C^{\S}$ | 13 (15.1) | 11 (19) | 2 (7.1) | 0.152 |
| Mortality 28 in days | 35 (40.7) | 26 (44.8) | 9 (32.1) | 0.262 |

SAPS III - Simplified Acute Physiology Score III; CRP - C-reactive protein. * p compares variables in relation to infusion volume $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg} ;{ }^{\dagger}$ well-nourished; ${ }^{\ddagger}$ risk of moderate undernutrition or malnutrition; ${ }^{\S}$ severe malnutrition. Results are expressed as the mean $\pm$ standard deviation (Student's $t$-test) or $n$ (\%) (chi-square test).

Table 2 - Presence of digestive tract disorders, over 5 days, considering all patients and, among them, those who received $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of body weight of fluid on the first day

| Variables | All | Infusion volume $>\mathbf{3 5 m L} / \mathbf{k g}$ |  | * p value |
| :---: | :---: | :---: | :---: | :---: |
|  | ( $\mathrm{n}=86$ ) | Yes ( $\mathrm{n}=58$ ) | No ( $\mathrm{n}=28$ ) |  |
| Constipation | 64 (74.4) | 42/58 (72.4\%) | 22/28 (78.6\%) | 0.854 |
| Diarrhea | 10 (11.6) | 7/58 (12\%) | 3/28 (10.7\%) | 0.540 |
| Abdominal distension | 20 (23.3) | 17/58 (29.3\%) | 3/28 (10.7\%) | 0.056 |
| Melena | 2 (2.3) | 2/58 (3.4\%) | 0/28 (0\%) | 0.320 |
| Drainage in NG tube (mL) | $505 \pm 412$ | $513 \pm 440$ | $460 \pm 251$ | 0.842 |

NG tube - nasogastric tube. * p compares variables in relation to infusion volume $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$. Results are expressed as n (\%) (chi-square test) or mean $\pm$ standard deviation Student's $t$-test).

Table 3 - Prescribed and infused enteral diet and protein-calorie deficiency, considering all patients and, among them, those who received $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$ of body weight of fluid on the first day

| Variables |  | Infusion volume $>35 \mathrm{~mL} / \mathrm{kg}$ |  | *p value |
| :---: | :---: | :---: | :---: | :---: |
|  | ( $\mathrm{n}=86$ ) | Yes ( $\mathrm{n}=58$ ) | No ( $\mathrm{n}=28$ ) |  |
| ENT prescribed 5 days (mL) | $4,297 \pm 1202$ | $4,163 \pm 1,245$ | $4,573 \pm 1,068$ | 0.140 |
| ENT infused 5 days (mL) | $2,994 \pm 1324$ | $2,781 \pm 1,338$ | $3,433 \pm 1,202$ | 0.032 |
| ENT prescribed on day 1 (mL) | $553 \pm 138$ | $547 \pm 139$ | $565 \pm 137$ | 0.577 |
| ENT infused on day 1 (mL) | $185 \pm 197$ | $179 \pm 205$ | $196 \pm 182$ | 0.717 |
| ENT infused 5 days (\%) | $67.8 \pm 19.8$ | $64.3 \pm 19.7$ | $74.9 \pm 18.6$ | 0.020 |
| Caloric deficit of 5 days (kcal) | $1813 \pm 850$ | $1868 \pm 870$ | $1699 \pm 811$ | 0.392 |
| Protein Deficit of 5 days (g) | $94.7 \pm 46$ | $95.9 \pm 46.3$ | $92.1 \pm 45.9$ | 0.722 |

ENT - enteral nutritional therapy. * p compares variables in relation to infusion volume $>35 \mathrm{~mL} / \mathrm{kg}$ or $\leq 35 \mathrm{~mL} / \mathrm{kg}$. Results are expressed as the mean $\pm$ standard deviation (Student's $t$-test).

Table 4 - Prescribed and infused enteral diet and protein-calorie deficiency, considering all patients and, among them, those that received sodium $\geq 8.73 \mathrm{~g}$ on the first day

| Variables | Sodium $\geq \mathbf{8 . 7 3 g}$ |  | *p value |
| :---: | :---: | :---: | :---: |
|  | Yes ( $\mathrm{n}=65$ ) | No ( $\mathrm{n}=21$ ) |  |
| ENT prescribed 5 days (mL) | $4,203 \pm 1,265$ | $4,586 \pm 948$ | 0.207 |
| ENT infused 5 days (mL) | $2,827 \pm 1,397$ | $3,509 \pm 911$ | 0.013 |
| ENT prescribed on day 1 (mL) | $540 \pm 134$ | $590 \pm 144$ | 0.149 |
| ENT infused on day 1 (mL) | $180 \pm 205$ | $198 \pm 175$ | 0.727 |
| ENT infused 5 days (\%) | $64.6 \pm 19.5$ | $77.6 \pm 17.9$ | 0.008 |
| Caloric deficit of 5 days (Kcal) | $1,903 \pm 879$ | 1,532 $\pm 700$ | 0.082 |
| Protein deficit of 5 days (g) | $100 \pm 49.3$ | $78 \pm 28.5$ | 0.014 |

ENT - enteral nutritional therapy. Results are expressed as the mean $\pm$ standard deviation (Student's $t$-test).
other nutritional variables studied ( $\mathrm{p}>0.05$ ). These results are provided in table 3 . The number of patients receiving ENT with volumes below $2,063 \mathrm{~mL}$ over 5 days (cutoff point, 25th percentile) was approximately 4.5 times higher in the group with the highest intravenous fluid infusion (19/58 (32.7\%) versus $2 / 28$ ( $7.1 \%$ ); $\mathrm{p}=0.010$ ).

## Sodium

Patients who received intravenous infusion $\geq 8.73 \mathrm{~g}$ sodium on the first day (cutoff point, 25th percentile) received a lower volume of enteral diet over 5 days ( 2,827
$\pm 1,397 \mathrm{~mL}$ versus $3.509 \pm 911.9 \mathrm{~mL} ; \mathrm{p}=0.013)$, had a lower percentage of infused diet ( $64.6 \pm 19.5 \%$ versus $77.6 \pm 17.9 \% ; \mathrm{p}=0.008$ ), and had a higher protein deficit ( $100 \pm 49.3$ versus $78 \pm 28.5 \mathrm{~g} ; \mathrm{p}=0.014$ ) than did those who received less than 8.73 g of sodium on the first day (Table 4).

## DISCUSSION

The findings showed that the administration of intravenous fluids on the first day of hospitalization at a volume greater than $35 \mathrm{~mL} / \mathrm{kg}$ may hinder enteral diet
infusion. In this study, the mean fluid administered was approximately $48 \mathrm{~mL} / \mathrm{kg}$, and some patients received almost $170 \mathrm{~mL} / \mathrm{kg}$ on the first day; some patients received almost 5 times the cutoff a mount of $35 \mathrm{~mL} / \mathrm{kg}$. The amount of sodium infused was also above the recommended cutoff amount, which is $2.0 \mathrm{~g} /$ day; ${ }^{(25)}$ some patients received 15 times this value. In addition to being rich in sodium, $0.9 \%$ saline solution is considered nonphysiological. ${ }^{(26)}$ According to Lobo et al., ${ }^{(16)}$ patients who received a volume of saline solution $\geq 3.0 \mathrm{~L} /$ day remained hospitalized longer and showed a delayed return of intestinal function, which may contribute to lower tolerance of enteral nutrition. Thus, although time is crucial in the resuscitation phase to achieve hemodynamic stability with the administration of fluids, in the first 3 hours, this overload can result in greater intestinal dysmotility. ${ }^{(27)}$ Positive fluid balance, intestinal loop edema, vomiting, gastroparesis and adynamic ileal are some of these complications. ${ }^{(11,12,14,15,28)}$ Alsous et al. ${ }^{(29)}$ showed that patients in septic shock with at least 1 day of negative water balance $>500 \mathrm{~mL}$ in the first 3 days had lower mortality. Excess fluid can also cause an increase in body weight, from 3.0 to 7.0 kg , which is associated with worse outcomes - mainly lower oxygen saturation and complications with surgical wounds. ${ }^{(30)}$ A recent study conducted with surgical patients showed a higher rate of infection at the surgical site and a higher risk of kidney injury, with no difference in septic complications and mortality between the restrictive group and liberal group. Although this study found benefits in the use of a more liberal volume, several others found results that favored patients who received a more restrictive volume of fluids. ${ }^{(31)}$ A study showed that the lower the intravenous fluid administration is, the greater the food intake. Patients who received less fluids (2.0L) and sodium ( 70 mmol ) per day had better gastric emptying and "willingness to feed", while the other group, which received a greater fluid volume, showed delayed gastric emptying and vomiting and did not feed normally. ${ }^{(28)}$ Additionally, in this study, patients from the restrictive group had earlier bowel recurrence and were discharged earlier. ${ }^{(28)}$ There is a close relationship between intestinal flow and the digestion and absorption of food; ${ }^{(16)}$ therefore, there are benefits of a more restrictive fluid and sodium protocol. Success in nutrition is related to lower fluid accumulation in the interstitium and greater weight loss due to increased fluid excretion, with consequent improvement of serum albumin. ${ }^{(32)}$ Thus, more restrictive fluid and sodium protocols may favor early oral/enteral
nutrition. ${ }^{(33)}$ The replacement and administration of sodium and fluids should be addressed with careful planning so that better clinical outcomes are achieved. ${ }^{(11)}$ In the resuscitation phase, stability may be achieved with a lower intravenous fluid load in the presence of vasoactive drugs. ${ }^{(22.23)}$ Our data showed that intravenous fluid and sodium overload may have contributed, in some way, to lower enteral diet infusion, which resulted in caloric and protein deficits. This deficit in the first days is not compensated in the days following the hospitalization of critically ill patients. ${ }^{(18)}$ Critical protein-calorie deficiency in the ICU is approximately $70 \%$ and is associated with a lower probability of accumulated survival. ${ }^{(34)}$ A study by our group showed that increased fluid administration is associated with lower enteral diet infusion and proteincalorie deficiency in patients in intensive care. ${ }^{(35)}$ Several other studies have shown that the reduced supply of calories and protein can increase infectious complications, hospitalization stays, and mortality. ${ }^{(36-38)}$

Another important context is tolerance to nutritional therapy and the digestive tract. Among the factors that interfere with diet tolerance are gastrointestinal disorders and diet composition, but these are not the only ones. ${ }^{(33,40)}$ Our data showed that infusion of intravenous fluids $>35 \mathrm{~mL} / \mathrm{kg}$ contributed to lower infusion of enteral nutrition over 5 days. In our study, we found no difference in the amount of norepinephrine administered or in the SAPS III score for patients who received more or less than $35 \mathrm{~mL} / \mathrm{kg}$. However, this study has a small number of cases, and thus, other multivariate analysis studies are needed to better study this relationship between the volume of intravenous fluids and the efficacy of enteral diet infusion in critically ill patients. Although this study has limitations, its findings may corroborate the development of more restrictive intravenous fluid administration protocols during the resuscitation phase. This more restrictive volume can result in better tolerance of an enteral diet as well as contribute to greater infusion of the diet with a subsequent reduction in calorie and protein deficits.

## CONCLUSION

The results of this study support the assumption that the administration of intravenous fluids, $>35 \mathrm{~mL} / \mathrm{kg}$ and sodium $\geq 8.73 \mathrm{~g}$, on the first day of hospitalization contributes to lower infusion of enteral diets in nonsurgical intensive care patients on mechanical ventilation.

## Authors' contributions

Suzana Souza Arantes participated in the study design, data collection and analysis, interpretation of the result, and writing of the article. José Eduardo de Aguilar-Nascimento participated in the data analysis and
writing of the article. João Manoel Silva Jr participated in the data analysis and writing and review of the article. Diana Borges Dock-Nascimento participated in the study design, data collection and analysis, interpretation of the data, and writing and review of the article.

## RESUMO

Objetivo: Avaliar os efeitos da administração intravenosa de fluidos e sódio no primeiro dia de internação com a infusão de nutrição enteral em pacientes de terapia intensiva.

Métodos: Estudo de coorte prospectivo realizado com pacientes críticos, não cirúrgicos, em ventilação mecânica internados pelo menos há 5 dias com nutrição enteral. Investigaram-se a quantidade de fluidos e sódio administrados por via venosa no primeiro dia e o volume de nutrição enteral infundido nos primeiros 5 dias. Comparou-se o volume de fluidos intravenosos do primeiro dia $>35 \mathrm{~mL} / \mathrm{kg}$ ou $\leq 35 \mathrm{~mL} / \mathrm{kg}$ de peso corporal e de sódio (acima ou abaixo do percentil 25), com o total de nutrição enteral infundida.

Resultados: Estudaram-se 86 pacientes com média ( $\pm$ desvio padrão) de $65 \pm 17$ anos, sendo $54,7 \%$ do sexo feminino. Foram administrados, no primeiro dia, $3.393,7 \pm 1.417,0 \mathrm{~mL}$ de
fluidos ( $48,2 \pm 23,0 \mathrm{~mL} / \mathrm{kg}$ ) e $12,2 \pm 5,1 \mathrm{~g}$ de sódio. Cinquenta e oito ( $67,4 \%$ ) pacientes receberam mais de $35 \mathrm{~mL} / \mathrm{kg}$ de fluidos. Em 5 dias, foram ofertados $67 \pm 19,8 \%(2.993,8 \pm 1.324,4 \mathrm{~mL})$ da nutrição enteral. Os pacientes que receberam $>35 \mathrm{~mL} / \mathrm{kg}$ de fluidos intravenosos também receberam menos nutrição enteral em 5 dias ( $2.781,4 \pm 1.337,9$ versus $3.433,6 \pm 1.202,2 \mathrm{~mL} ; \mathrm{p}=$ 0,03 ) versus quem recebeu $\leq 35 \mathrm{~mL} / \mathrm{kg}$. Pacientes com infusão de sódio intravenoso acima do percentil $25(\geq 8,73 \mathrm{~g})$ no primeiro dia receberam menos volume de nutrição enteral em 5 dias $(2.827,2 \pm 1.398,0$ versus $3.509,3 \pm 911,9 \mathrm{~mL} ; \mathrm{p}=0,02)$.

Conclusáo: Os resultados deste estudo apoiam o pressuposto de que a administração de fluidos intravenosos no primeiro dia de internação $>35 \mathrm{~mL} / \mathrm{kg}$ e de sódio $\geq 8,73 \mathrm{~g}$ pode contribuir para a menor infusão de nutrição enteral em pacientes críticos.

Descritores: Cuidados intensivos; Fluido; Sódio; Nutrição enteral

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