



OPEN ACCESS

EDITED AND REVIEWED BY
Paula Ravasco,
Catholic University of Portugal, Portugal

*CORRESPONDENCE
Cristian Deana
✉ deana.cristian@gmail.com

RECEIVED 16 September 2023
ACCEPTED 22 September 2023
PUBLISHED 16 October 2023

CITATION
Deana C, Vecchiarelli P, Picetti E and Molino A
(2023) Editorial: Intermittent feeding in critically
ill patients. *Front. Nutr.* 10:1295405.
doi: 10.3389/fnut.2023.1295405

COPYRIGHT
© 2023 Deana, Vecchiarelli, Picetti and Molino.
This is an open-access article distributed under
the terms of the [Creative Commons Attribution
License \(CC BY\)](https://creativecommons.org/licenses/by/4.0/). The use, distribution or
reproduction in other forums is permitted,
provided the original author(s) and the
copyright owner(s) are credited and that the
original publication in this journal is cited, in
accordance with accepted academic practice.
No use, distribution or reproduction is
permitted which does not comply with these
terms.

Editorial: Intermittent feeding in critically ill patients

Cristian Deana^{1*}, Pietro Vecchiarelli², Edoardo Picetti³ and Alessio Molino⁴

¹Department of Anesthesia and Intensive Care, Health Integrated Agency of Friuli Centrale, Udine, Italy, ²Intensive Care Unit, Ospedale Belcolle, Viterbo, Italy, ³Department of Anesthesia and Intensive Care, Parma University Hospital, Parma, Italy, ⁴Department of Translational and Precision Medicine, Sapienza University of Rome, Rome, Italy

KEYWORDS

enteral nutrition, critically ill patient, parenteral nutrition, intermittent feeding, autophagia, refeeding syndrome, proteins

Editorial on the Research Topic

Intermittent feeding in critically ill patients

Critical illness deeply alters the patients' metabolism, determining a high catabolic state since the early phase of the intensive care unit (ICU) stay. Thus, the risk to develop toward malnutrition is high especially if an adequate artificial nutrition therapy is not provided (1).

However, we are far from having available a unique nutritional prescription able to "fit" all in an ICU setting. We should consider that artificial nutrition may confer benefit to the patients, but also could be harmful, if not adequately and timely delivered.

One of the main aspects that makes artificial nutrition complex in ICU is represented by the heterogeneous population admitted for the different acute diseases. Sir Cuthbertson in the 50's described different phases of critically illness, with a non-inhabitable catabolic state at the initial phase of the disease, then becoming anabolic only when the storm has passed. Unfortunately, we have no precise markers for the transition from one phase to another.

For this reason, clinical trials that started early full nutrition in order to reduce the catabolism failed to provide clinical benefits. Indeed, researchers observed that an excess of calories and proteins too early in less severe ICU patients may increase mortality rates (2).

In this regard, the amount of proteins plays a key role during the early phase of critically ill patients: some large randomized trials demonstrated no advantage on mortality with higher compared with lower protein nutrition content (3). Furthermore, evidences suggest that high protein load administered too early could impair autophagy, a protective mechanism that removes damaged cells (4).

The route by which artificial nutrition is delivered is another important point. The use of early parenteral nutrition (PN) was revisited after the EPANIC trial in whom patients that received late (after 7 days from ICU admission) PN had faster recovery, fewer complications and indirectly produced lower health related costs (5).

However, when PN has to be started, for example when enteral route is contraindicated, some advices have to be considered to reduce potential complications, such as catheter-related blood stream infections and thrombosis (Zaccone et al.; Ko et al.).

Enteral nutrition is nowadays the preferred route of feeding critically ill patients, contributing to maintain a trophic intestinal mucosa. Continuous enteral feeding is widely used because it is easier and require less effort for personnel.

However, it does not reproduce the physiologic response to muscle protein synthesis after protein bolus. Moreover, continuous tube feeding abolishes the entero-hormonal

response. Intermittent feeding could be better compared to continuous feeding considering that it can enhance muscular protein synthesis, and the fasting periods may be protective through enhanced autophagy and ketogenesis (6).

Moreover, both methods do not prevent gastrointestinal complications, such as diarrhea or constipation [(7), Qu et al.].

An important aspect to consider when initiating artificial nutrition after a prolonged fasting state is the refeeding syndrome. Low phosphate levels are the main characteristic of this complication, but other electrolytes disturbances should be taken into consideration.

In neurocritically ill patients, often necessitating enteral nutrition due to swallowing disturbances, some risk factors, among them APACHE II score, SOFA score, low serum albumin and low circulating levels of potassium, have been indicated by Zhang et al..

Sepsis is another severe condition where deep disturbances in metabolism are often observed. Evidences pointed out that vitamin C plus hydrocortisone and thiamine reduced mortality in septic patients (8). However, Liang et al. in their meta-analysis did not confirm this aspect, suggesting new studies investigating this important topic.

Guan et al. in a retrospective study including more than 19,000 patients, found that vitamin D supplementation in ICU patients reduced the risk for sepsis and new mechanical ventilation, but not mortality at 28-day.

Based on several evidences, an ideal nutrition is far from actual practices. However, moving toward precise and personalized medicine also among ICU patients, probably using in the next years artificial intelligence models, represents an intriguing scenario.

Also, the continuum of care, from the hospital to the rehabilitation units and finally in a home setting, should be effective

in reducing the incidence of malnutrition with its negative long-term effects after a critical illness [(9); Diamanti et al.].

Author contributions

CD: Writing—original draft, Writing—review and editing. PV: Writing—original draft, Writing—review and editing. EP: Writing—original draft, Writing—review and editing. AM: Writing—original draft, Writing—review and editing.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

The author(s) declared that they were an editorial board member of Frontiers, at the time of submission. This had no impact on the peer review process and the final decision.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

References

- Preiser JC, Ichai C, Orban JC, Groeneveld AB. Metabolic response to the stress of critical illness. *Br J Anaesth.* (2014) 113:945–54. doi: 10.1093/bja/aeu187
- Pardo E, Lescot T, Preiser JC, Massanet P, Pons A, Jaber S, et al. Association between early nutrition support and 28-day mortality in critically ill patients: the FRANS prospective nutrition cohort study. *Crit Care.* (2023) 27:7. doi: 10.1186/s13054-022-04298-1
- Heyland DK, Patel J, Compher C, Rice TW, Bear DE, Lee ZY, et al. The effect of higher protein dosing in critically ill patients with high nutritional risk (EFFORT Protein): an international, multicentre, pragmatic, registry-based randomized trial. *Lancet.* (2023) 401:568–76. doi: 10.1016/S0140-6736(22)02469-2
- Casaer MP, Wilmer A, Hermans G, Wouters PJ, Mesotten D, Van den Berghe G. Role of disease and macronutrient dose in the randomized controlled EPaNIC trial: a *post-hoc* analysis. *Am J Respir Crit Care Med.* (2013) 187:247–55. doi: 10.1164/rccm.201206-0999OC
- Casaer MP, Mesotten D, Hermans G, Wouters PJ, Schetz M, Meyfroidt G, et al. Early vs. late parenteral nutrition in critically ill adults. *N Engl J Med.* (2011) 365:506–17. doi: 10.1056/NEJMoa1102662
- Bear DE, Hart N, Puthucherry Z. Continuous or intermittent feeding: pros and cons. *Curr Opin Crit Care.* (2018) 24:256–61. doi: 10.1097/MCC.0000000000000513
- Danielis M, Mattiussi E, Piani T, Iacobucci A, Tullio A, Molino AV, et al. Diarrhea and constipation during artificial nutrition in intensive care unit: a prospective observational study. *Clin Nutri.* 4:7. doi: 10.1016/j.clnesp.2023.07.007
- Marik PE, Khangoora V, Rivera R, Hooper MH, Catravas J. Hydrocortisone, Vitamin C, and Thiamine for the treatment of severe sepsis and septic shock: a retrospective before-after study. *Chest.* (2017) 151:1229–38. doi: 10.1016/j.chest.2016.11.036
- Deana C, Vetrugno L, Cortegiani A, Mongodi S, Salve G, Mangiagalli M, et al. Quality of life in COVID-related ARDS patients 1-year after intensive care discharge (Odissea study): a multicenter observational study. *J Clin Med.* (2023) 12:1058. doi: 10.3390/jcm12031058