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KABI**

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United For Clinical Nutrition EuroPN Survey¹

Medical nutrition therapy
and clinical outcomes in
critically ill adults:

A European multinational,
prospective observational
cohort study



1. Matejovic, M, Huet, O, Dams, K, Elke G, Vaquerizo Alonso C, Csomos A, Krzych LJ, Tetamo R, Puthucheary Z, Rooyackers O, Tjäder I, Küchenhoff H, Hartl WH, Hiesmayr M. Medical nutrition therapy and clinical outcomes in critically ill adults: a European multinational, prospective observational cohort study (EuroPN). *Critical Care*. 2022;26(143).



Background

Why research in clinical nutrition in the ICU matters

Adequate nutrition is essential to support the recovery of hospitalized patients, particularly those with critical illness.^{2,3}



ESPEN ICU guidelines recommend a medical nutrition therapy that:

- provides <70% of measured energy expenditure (EE) or of estimated needs during the early phase of acute illness⁴
- provides 80-100% of EE after day 3⁴
- enables the progressive delivery of 1.3 g/kg protein equivalents⁴

If IC is not used, one of these options may be preferred:

- VO_2 or VCO_2 measurements
- simple weight-based equations (such as 20-25 kcal/kg/d)⁴



Background

Why research in clinical nutrition in the ICU matters



Addressing unmet needs



- Most critically ill patients do not receive adequate nutritional intake according to guideline targets.^{5,6}
- Evidence from RCTs and observational studies on the amount and timing of medical nutrition therapy and clinical outcomes is inconsistent.⁷⁻¹⁵



Study design*

Exploring how medical nutrition is put into practice across Europe

EuroPN Survey: One of the largest longitudinal European real-world-studies

- **Multinational:**

- Austria
- Belgium
- Czech Republic
- France
- Germany
- Hungary
- Italy
- Poland
- Spain
- Sweden
- UK

- **Prospective, observational**

- Data relating to nutrition intake was collected in the ICU for up to **15 days** and compared to the **ESPEN guidelines targets**

- **77 ICUs, 1172 patients**

- **Follow-up until day 90**



* Full study protocol published: Hiesmayr M, et al. 2021.¹⁶

ESPEN: European Society for Clinical Nutrition and Metabolism,
ICU: Intensive care unit



Study design*

Exploring how medical nutrition is put into practice across Europe

Study population

Patient Characteristics (n=1172 ^a)	Number of (%) or Median [95% CI]
Age	66.0 [56.0;74.0]
Male	745 (63.6)
Female	427 (36.4)
BMI (kg/m ²), n=1168	26.8 [24.0;31.1]
Type of ICU admission	
Surgical (emergency/elective)	580 (49.5)
Non-surgical	573 (48.9)
Others	19 (1.6)
APACHE II, n=1132	18.5 [13.0;26.0]
SOFA (Median), n=1042	7 [4;10]
ICU LOS (days), n=1158	10 [7;16]

^a unless otherwise indicated

Aims

- Describe medical nutrition therapy for up to 15 days after ICU admission in European critically ill patients with a minimum length of stay of 5 days
- Assess the association between the daily calorie and protein intake and time to weaning from invasive mechanical ventilation and 90-day survival time



APACHE II Score: Acute Physiology And Chronic Health Evaluation, BMI: Body mass index, CI: Confidence interval, EN: Enteral nutrition, ESPEN: European Society for Clinical Nutrition and Metabolism, ICU: Intensive care unit, LOS: Length of stay, SOFA: Sequential Organ Failure Assessment

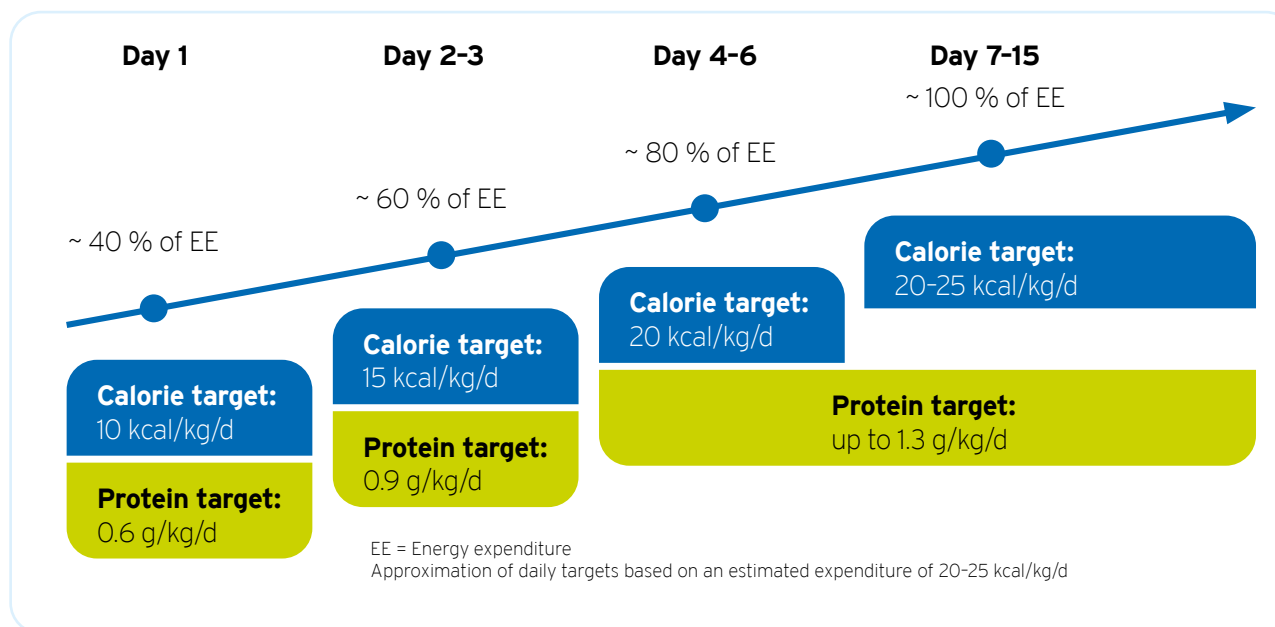
* Full study protocol published: Hiesmayr M, et al. 2021.¹⁶



Methods

Connecting nutritional intake with nutritional targets and clinical outcomes

Calculation of calorie and protein balances* based on progressive daily targets



* Calculated as the percentage deviation of nutritional intakes from all sources compared to ESPEN targets

EE: Energy expenditure, ESPEN: European Society for Clinical Nutrition and Metabolism



Methods

Connecting nutritional intake with nutritional targets and clinical outcomes

Association of daily nutrition intake and outcomes

Daily nutrient intake	LOW	MODERATE	HIGH
Calories (kcal/kg/d)	< 10	10-20	> 20
Protein (g/kg/d)	< 0.8	0.8-1.2	> 1.2



Clinical Outcome

- 90-day survival time
- Time to successful weaning from invasive mechanical ventilation (IMV)



Results

How well are targets being met?

Valuable insights into the current situation.



Medical nutrition therapy

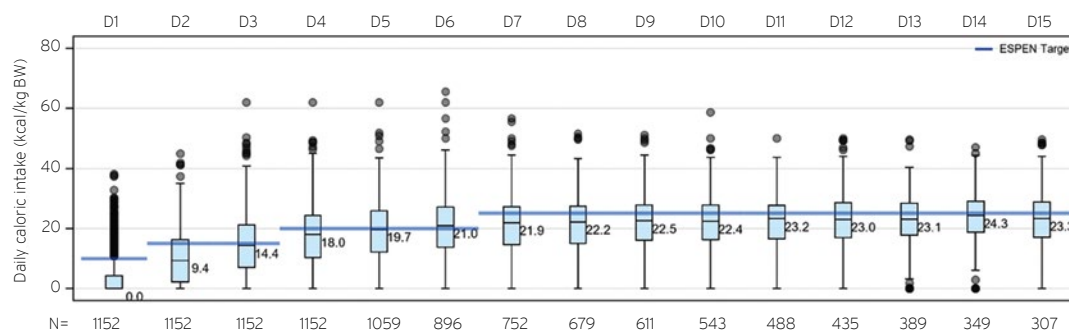
- EN and PN were initiated on median ICU day 2.0 [95% CI: 2.0;4.0].
The provision of nutrition was increased progressively over the first 5 days.
- Over half of the patients received medical nutrition therapy, either EN and/or PN, on day 3 after ICU admission.



Results

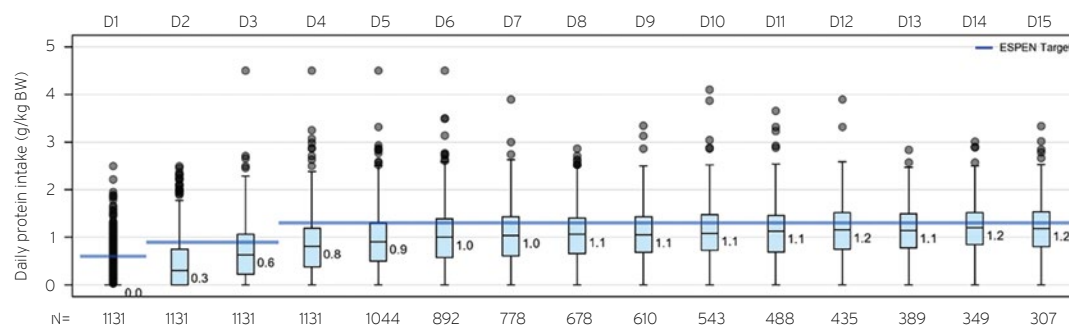
How well are targets being met?
Valuable insights into the current situation.

Total daily caloric intake vs. daily ESPEN target[†]



83% (Median, 95% CI 59.2;106.6)
of patients' ESPEN calorie targets were met

Total daily protein intake vs. daily ESPEN target[†]



65% (Median, 95% CI 41.4;90.9)
of patients' ESPEN protein targets were met

[†]Intake is presented as the median value, interquartile range, minimum and maximum values with outliers versus pre-defined targets (blue horizontal bars) based on the 2019 ESPEN Guideline on clinical nutrition in the ICU.⁴

BW: Bodyweight, CI: Confidence interval, EN: Enteral nutrition, ESPEN: European Society for Clinical Nutrition and Metabolism, ICU: Intensive care unit, PN: Parenteral nutrition



Clinical outcomes

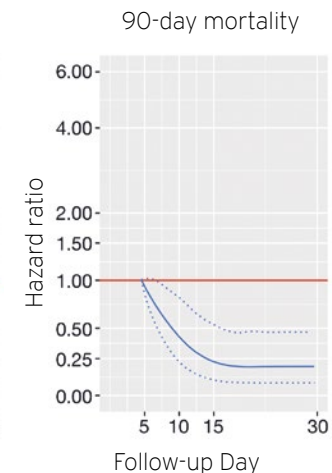
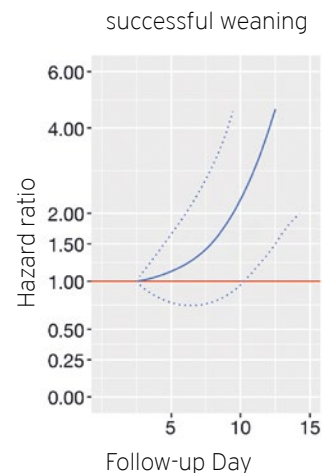
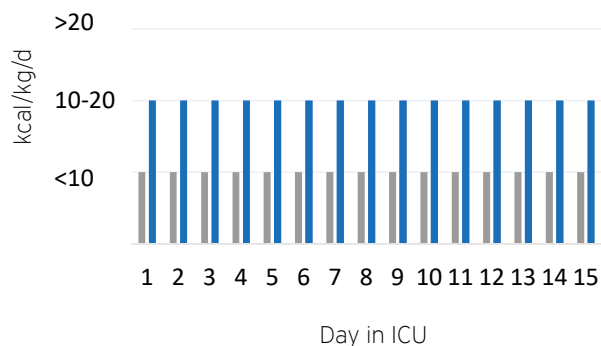
How do patients benefit from moderate medical nutrition?



Calories[†]

10-20 kcal/kg/day (moderate) from day 1 to day 15 was significantly associated with a **longer 90-day survival time and shorter time on IMV** compared to less calories

Comparison of hypothetical calorie intakes (low vs. moderate)



[†] Hypothetical medical nutrition therapies with different levels of daily calorie and protein intake were used to facilitate the time-varying hazard ratios (HR) between nutrition and outcomes. Reference medical nutrition therapy is thereby always the one providing fewer calories. Blue solid lines indicate the Hazard ratio (HR), hatched lines the corresponding 95% CI; Significant association was indicated when HR and 95% CI do not cross the red line. A HR and 95% CI < 1 would indicate a longer survival time but also a longer time until extubation associated with the medical nutrition therapy providing more calories.

CI: Confidence interval, HR: Hazard ratio, ICU: Intensive care unit, IMV: Invasive mechanical ventilation



Clinical outcomes

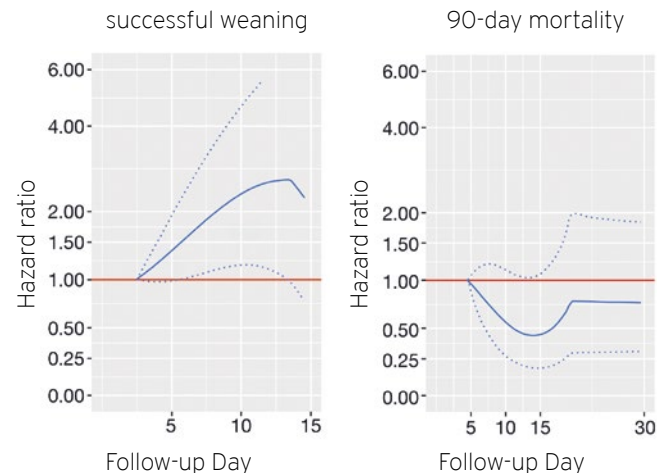
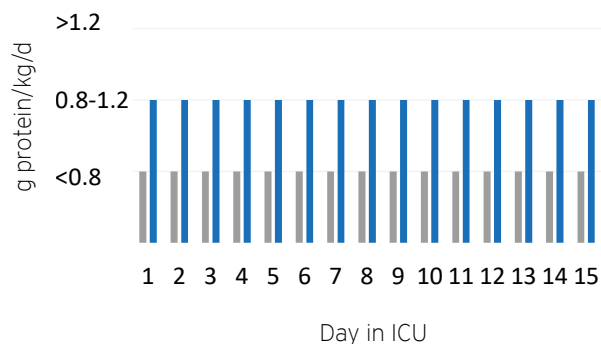
How do patients benefit from moderate medical nutrition?



Proteins[†]

0.8-1.2 g/kg/day (moderate) from day 1 to day 15 was significantly associated with **earlier weaning from IMV**, but not with survival, compared to less protein

Comparison of hypothetical protein intakes (low vs. moderate)



[†] Hypothetical medical nutrition therapies with different levels of daily calorie and protein intake were used to facilitate the time-varying hazard ratios (HR) between nutrition and outcomes. Reference medical nutrition therapy is thereby always the one providing fewer calories. Blue solid lines indicate the Hazard ratio (HR), hatched lines the corresponding 95% CI; Significant association was indicated when HR and 95% CI do not cross the red line. A HR and 95% CI < 1 would indicate a longer survival time but also a longer time until extubation associated with the medical nutrition therapy providing more calories.

CI: Confidence interval, HR: Hazard ratio, ICU: Intensive care unit, IMV: Invasive mechanical ventilation



Conclusion

New evidence on the importance of medical nutrition therapy



Main take-aways of the study

- Calorie intake was mainly provided according to the targets recommended by the ESPEN guideline, but protein intake was lower.
- Early moderate daily calorie and protein intake were associated with improved clinical outcomes.

United For Clinical Nutrition EuroPN Survey:

The EuroPN study adds to the existing evidence on the benefits of an early medical nutrition therapy with moderate calorie and protein intake on clinical outcomes in critically ill patients.





Join us in raising the relevance of clinical nutrition!



Learn more about the initiative and explore the results of our multinational **United for Clinical Nutrition EuroPN study** on <https://eu.unitedforclinicalnutrition.com>

References

- Matejovic, M, Huet, O, Dams, K, et al. Medical nutrition therapy and clinical outcomes in critically ill adults: a European multinational, prospective observational cohort study (EuroPN). *Critical Care*. 2022;26(143). <https://doi.org/10.1186/s13054-022-03997-z>
- Stoppe C, van Gassel R, Jonckheer J, et al. First international meeting of early career investigators: Current opportunities, challenges and horizon in critical care nutrition research. *Clinical Nutrition ESPEN*. 2020;40:92-100.
- Puthuchery ZA, Rawal J, McPhail M, et al. Acute Skeletal Muscle Wasting in Critical Illness. *JAMA*. 2013;310(15):1591-1600.
- Singer P, Blaser AR, Berger MM, et al. ESPEN guideline on clinical nutrition in the intensive care unit. *Clinical Nutrition*. 2019;38(1):48-79.
- Heyland DK, Dhaliwal R, Wang M, et al. The prevalence of iatrogenic underfeeding in the nutritionally 'at-risk/critically ill patient: Results of an international, multicenter, prospective study. *Clinical Nutrition*. 2015;34(4):659-66.
- Vallejo KP, Martínez CM, Adames AAM, et al. Current clinical nutrition practices in critically ill patients in Latin America: a multinational observational study. *Critical Care*. 2017;21(1):1-11.
- Arabi YM, Aldawood AS, Haddad SH, et al. Permissive under-feeding or standard enteral feeding in critically ill adults. *New England Journal of Medicine*. 2015;372(25):2398-408.
- Zusman O, Theilla M, Cohen J, Kagan I, et al. Resting energy expenditure, calorie and protein consumption in critically ill patients: a retrospective cohort study. *Critical Care*. 2016;20(1):1-8.
- Bendavid I, Zusman O, Kagan I, et al. Early administration of protein in critically ill patients: a retrospective cohort study. *Nutrients*. 2019;11(1):106.
- Singer P, Anbar R, Cohen J, et al. The tight calorie control study (TICACOS): a prospective, randomized, controlled pilot study of nutritional support in critically ill patients. *Intensive Care Medicine*. 2011;37(4):601-9.
- Heidegger CP, Berger MM, Graf S, et al. Optimisation of energy provision with supplemental parenteral nutrition in critically ill patients: a randomised controlled clinical trial. *The Lancet*. 2013;381(9864):385-93.
- Weijs PJ, Stapel SN, de Groot SD, et al. Optimal protein and energy nutrition decreases mortality in mechanically ventilated, critically ill patients: a prospective observational cohort study. *Journal of Parenteral and Enteral Nutrition*. 2012;36(1):60-8.
- Compher C, Chittams J, Sammarco T, et al. Greater protein and energy intake may be associated with improved mortality in higher risk critically ill patients: a multicenter, multinational observational study. *Critical Care Medicine*. 2017;45(2):156-63.
- Allingstrup MJ, Kondrup J, Wiis J, et al. Early goal-directed nutrition versus standard of care in adult intensive care patients: the single-centre, randomised, outcome assessor-blinded EAT-ICU trial. *Intensive Care Medicine*. 2017;43(11):1637-47.
- Koekkoek WACK, van Setten CHC, Olthof LE, et al. Timing of PROTEIN Intake and clinical outcomes of adult critically ill patients on prolonged mechanical VENTilation: The PROTINVENT retrospective study. *Clinical Nutrition*. 2019;38(2):883-90
- Hiesmayr M, Csomos A, Dams K et al. Protocol for a prospective cohort study on the use of clinical nutrition and assessment of long-term clinical and functional outcomes in critically ill adult patients. *Clinical Nutrition ESPEN*. 2021;43:104-10.



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