Sepsis in Critical Care

One Sepsis Fits All? Are There Different Phenotypes of Sepsis? Diagnostic Approaches and Therapies, A. Edel, S. J. Schaller

Sepsis in Critical Care: Effective Antimicrobial Strategies in ICU, G. B. Nair, M. S. Niederman

The Alphabet Book of Sepsis, M. Leone


Sepsis Surveillance (Sepsis Sniffer): Where We Are Now and Where We Are Going, Y. Pinevich, B. W. Pickering, V. Herasevich

Symmetrical Peripheral Gangrene, C. B. Noel, J. L. Bartock, P. Dellinger


Understanding Carbon Dioxide in Resuscitation F. S. Zimmerman, G. Pachys, E. A. Alpert, S. Einav
ICU Management of the Very Old: The Evidence Base Anno 2022

This overview explores the publications concerning the very old ICU patients since 2011 and with a focus on publications from the VIP network on elderly COVID-19 patients.

Introduction.
In 2017, an international group of intensivists with a particular interest for geriatric intensive care published a statement paper (Flaatten et al. 2017). This paper discussed the status of research in the very old critical ill patients, and what should be the priorities for such research in the coming years (Table 1). In this overview we seek to find out where this area of research stands five years later. Are any of the suggested topics better described and have new and important issues emerged?

If we look at the absolute numbers of related clinical research in this group, we see a considerable increase from 2011 until today, in total 2124 publications, but the number of publications has stabilised the last five years around 300 publications per year (Figure 1). Hence publications on the very old ICU patients measured as published paper seems still to attain a high interest.

The topics of the publications are more diverse. With the same research string but adding 11 specific topics found either in the title or the abstract, we found a more diverse picture. The topics of interest spread from low interest like severity score (n=20) to high interest like outcome (n=485) (Figure 2). One topic occurred and peaked in 2020 and 2021 - COVID-19. We found this despite only covering two years to be overall the third most popular topic over the 5-year period and was by far the highest in 2021 with 142 papers. The topics within the COVID-19 subgroup naturally varied to cover a broad spectrum.

Comparing these findings with Table 1 reveals that several items found to be important in 2017 are poorly covered. This goes for the more soft items like the study of opinions among elderly about intensive care, end-of life issues, frailty and sedation. These are all important topics that merit more focus.

COVID-19 and the Elderly Group of ICU Patients
Old patients have paid a severe tribute to COVID with a disproportionate number needing admission to an intensive care and a large proportion succumbing from the disease. In the beginning of the pandemic, several countries issued recommendations for admission and treatment of the most severe cases. However, little was known about the validity and applicability of such recommendations for the group of old critically ill patients. Since most of the interventional studies excluded old patients or included only a small number, there is doubt for extrapolating data to an old population.

Our VIP network has been active since 2017 and it was straightforward to adapt our CRF to the COVID surges and to motivate centres to participate. As a result, the COVID-19 in very old ICU patients (COVIP) study recruited 3140 patients older than 70 years from 19 March 2020 to 4 February 2021 in more than 150 ICUs across 15 European countries. COVIP is a prospective observational study looking at
patient demographics, treatment modalities in ICU and outcomes included health related quality of life up to three months after admission to the ICU.

We confirmed the poor prognosis in this old population with only 39% of the patients surviving up to 90-days (Jung et al. 2021a). Further, we documented that the effect of crude age is less important that the degree of frailty in this group. Outcome in patients above 70 in the presence of frailty was not influenced by age. In our quality analysis of outcome within the group of surviving patients 48% (n = 592) experienced ‘severe problems’ or ‘extreme problems’ in at least one of the five domains of the EQ-5D-5L questionnaire (Soliman et al. 2022). ‘Severe problems’ were mentioned by 41% (n = 496), and ‘extreme problems’ on one of the five domains in 30% (n = 371).

Another important issue included in the COVIP study was an analysis of the use of steroids. It was commonly acknowledged that systemic steroids were beneficial for hospitalised patients with COVID-19, but with insufficient documentation to conclude in the very old patient group. In a comparison of patients receiving steroids or not during the first and second surge in Europe, we found an independent association of steroids with increased mortality from 14 days after ICU admission (Jung et al. 2021b).

Early tracheostomy was not found to be of benefit, and we documented huge variability in the use of this procedure across European countries (Polok et al. 2021). We were expecting improved prognosis from severe COVID-19 from the first to the second surge in parallel with optimisation of oxygenation therapy, avoidance of early invasive mechanical ventilation and use of steroids. In fact, we found treatments to differ during the second wave with less invasive mechanical ventilation and more use of steroids. However, these differences did not translate into better outcomes since the mortality was significantly higher in the second wave compared to the first one (Jung et al. 2021c). This higher mortality could be related to different admission policy and pressure on ICU beds (Jung et al. 2021c) and to the above mentioned use of steroid treatment that was detrimental in old patients.

We have further shown that pressure on ICU beds increased the number of decisions to limit life-sustaining treatment (LST) (Jung et al. 2022). We will claim that information on withholding and withdrawing LST is crucial to interpret survival curves and have been missing in most outcome papers including critical ill elderly patients (Flaatten et al. 2022).

We also took advantage of information from a previous cohort in patients without COVID (VIP2 study) to compare patient characteristics, treatment and outcome between COVID and non-COVID patients older than 80 years admitted for acute respiratory failure (Guidet et al. 2022). This was obtained with matching the propensity score and regression analysis patients from the two cohorts. In this study

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<th>TOPIC</th>
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<td>2</td>
<td>What is the opinion of octogenarians towards use of critical care resources in acute, severe vital organ failure? A European survey among 10,000 octogenarians</td>
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<td>The effects of including a geriatrician in the early assessment and discharge of octogenarians</td>
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<td>The effects of non-pharmacological interventions to reduce delirium in the ICU</td>
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Table 1. Priorities for research in very old critical ill patients
Very Old ICU Patients

We found elderly COVIP patients to be less sick (lower SOFA score), less frail (lower CFS) but with more decision to forgo LST and with a higher one-month mortality.

Conclusions
This overview reveals an increase in publications about critically ill ICU patients during the last five years. However, the topics of interest for publications follow traditional patterns with outcome, sepsis, and mechanical ventilation as the most popular issues. An exception is the large number of publications following the COVID-19 pandemic. Specific age-related issues like frailty, end-of-life issues and prognostications (severity score) are less frequently published. We could not find any publications related to the wishes of elderly persons or their family with regards to receiving intensive care.

We hope our VIP network will continue to reveal new knowledge about the very old ICU patients also in the future. More knowledge of this group is crucial if we want to overcome the challenge of the rapid increase in the population of elderly across the globe, and concomitant huge increase in very old ICU patients. We should not stop to treat elderly critical ill patients, but we need to know more about those who will not profit from such intervention for the benefit of the elderly themselves and their families.

Conflict of interest
None.

References
Flaatten H et al. (2017) The status of intensive care medicine research and a future agenda for very old patients in the ICU. Int Care Med. 43 1–10.

Figure 2. Major topics in the studies of the very old intensive care patients from 2017 to 2021