COVID-19 AND POST INTENSIVE CARE SYNDROME: A CALL FOR ACTION

Henk J. STAM, MD, PhD¹, Gerold STUCKI, MD, MS^{2,3}, Jerome BICKENBACH, LLB, PhD^{2,3} on behalf of the European Academy of Rehabilitation Medicine

From the ¹Department of Rehabilitation Medicine, Erasmus University Medical Center, Rotterdam, The Netherlands, ²Center for Rehabilitation in Global Health Systems, WHO Collaborating Center, Department of Health Sciences and Medicine, University of Lucerne, Switzerland and ³Swiss Paraplegic Research (SPF), Nottwil, Switzerland

Although we are currently overwhelmed by the astonishing speed of infection of the Covid-19 pandemic, and the daily onslaught of new, and ever-worsening predictions, it is vital that we begin to prepare for the aftershocks of the pandemic. Prominent among this will be the cohort of post-intensive case survivors who have been mechanically ventilated and will likely experience short- and medium-term consequences. The notion that patients surviving intensive care and mechanical ventilation for several weeks can be discharged home without further medical attention is a dangerous illusion. Post Intensive Care Syndrome and other severe conditions will require not only adequate screening but early rehabilitation and other interventions. Action must be taken now to prepare for this inevitable aftershock to the healthcare system.

Key words: Covid-19; Post Intensive Care Syndrome; Rehabilitation.

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Correspondence address: Gerold Stucki, Center for Rehabilitation in Global Health Systems, WHO Collaborating Center, Department of Health Sciences and Medicine, University of Lucerne, Switzerland

In 2020, the world community is experiencing a pandemic from a novel coronavirus Covid-19 that touches every corner of the world. During a pandemic, it is easy to get overwhelmed by the astonishing speed of infection and the daily onslaught of new, and everworsening predictions. Pandemics are extraordinary public health challenges that demand targeted and coordinated responses by health care systems and governments. But this pandemic, almost from the outset, has also raised the specter of utter health systems failures to cope with the 'surge' of demand, putting not only Covid-19 victims at risk but all other individuals who require health care services.

Researchers at the Institute of Health Metrics and Evaluation (IHME) at the University of Washington in Seattle, Washington, USA, modeled expected impacts of the pandemic on healthcare resources (1). They estimated that by mid-May the excess demand from Covid-19 is predicted to be nearly 65,000 hospital beds, and 17,000 intensive care unit (ICU) beds, while ventilator use is predicted to be as high as 20,000. These data constitute a burden that "is well beyond the current capacity of hospitals to manage, especially for ICU care" (2). It is expected that the results for the rest of the world, once they are available, will be worse, especially for low and middle resource countries (3).

It is not premature to begin to think past the pandemic when incidence and death rates are reduced to a point where more general population immunity – enhanced by the eventual vaccination options – becomes likely. From our current perspective, it is difficult to imagine this, but we must not only keep this in mind but urgently prepare for what amounts to the more long-lasting aftershocks of the pandemic.

As disease-causing respiratory distress, Covid-19 in a minority of cases (the Italian experience suggests around 15%) (4) often involving elderly individuals or those with chronic comorbidities, requires intensive care and reliance on ventilators. A distinctive feature of this disease is that when necessary, acute and intensive care and ventilator reliance are required for a considerably longer period – perhaps as high as 20 days (5, 6) – than more typical uses of ICU. Until countries begin to experience a decrease in incidence, these hospitalization and ICU usage rates will be a major concern. But as patients improve and are discharged home, the pandemic will cause an aftershock: recovery from the consequences of severe respiratory illness and the secondary disabilities that result from intensive care treatments, including Critical Illness Polyneuropathy (CIP) and Critical Illness Myopathy (CIM), as part of the Post Intensive Care Syndrome (PICS) (Video¹; Table I) (7).

¹https://youtu.be/FiU2we_9f1s

Table I. Post Intensive Care Syndrome (PICS)

Cognitive impairments Memory Attention Visuo-spatial Psychomotor Impulsivity Psvchiatric illness Anxiety Depression Post traumatic stress disorder Physical impairments Dyspnea/Impaired pulmonary function Pain Sexual dysfunction Impaired exercise tolerance Neuropathies Muscle weakness/Paresis Severe fatigue

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The notion that patients surviving intensive care and mechanical ventilation for several weeks can be discharged home without further medical attention is a dangerous illusion. The longer a patient remains in the ICU, the higher the risk for long-term physical, cognitive and emotional complications. Among the major risk factors are cognitive impairments, acute brain dysfunction, hypoxia, hypotension, and glucose dysregulation (8). Cognitive impairments, including delirium, have been reported in between 30% and 80% of post-ICU patients, varying in severity and duration depending on pre-existing cognitive deficits and age (9). The risk factors for PICS are shown in Table II. One-year outcomes of PICS have been reported to include new symptoms such as dyspnea, pain, sexual dysfunction, impaired pulmonary function and impaired exercise tolerance. But the most common complications are neuromuscular, resulting in poor mobility, frequent falls, even quadriparesis (6). CIP and CIM syndromes seen in approximately 25-45% of critically ill patients during and after intensive care stays with mechanical ventilation - exhibit even more severe neurodegenerative complications, including flaccid and symmetric paralysis, limb and respiratory muscle weakness, systemic inflammatory response syndrome, or multiple organ failure (10).

The long-term impacts of extended periods in ICU are not only highly disruptive to the health of individuals; but also to the society (11–13). Nearly a third of patients who experience PICS do not go back to work, and another third do not go back to their pre-ICU job, or a job with a pre-ICU salary. At least 25% of these patients experience a dramatic loss of independence and require assistance in activities in daily living one year after ICU admission, putting an enormous burden on informal and formal caregivers (14). Patients experiencing CIP suffer from decreased exercise capacity, disability and compromised quality of life for months, even years after intensive care.

There is a range of established clinical responses to these syndromes that control risk factors to avoid or limit the use of neuromuscular blocking agents and corticosteroids (15,16). There is a clear consensus that "early rehabilitation combining mobilization with

Table II. Risk	Factors for	- Post	Intensive	Care	Syndrome	(PICS)
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Delirium Duration of ICU admission Duration of sedation Duration of mechanical ventilation Age Hypoxia and hypotension Sepsis Glucose dysregulation Premorbid mental and physical comorbidity ICU: intensive care unit. physiotherapy is emerging as an important strategy to treat critical illness polyneuropathy and myopathy, and to facilitate and improve long-term recovery and functional independence of patients, and shorten the duration of ventilation and hospitalization" (8).

CALL FOR ACTION

Now is the time to acknowledge and take seriously the need for a coordinated response to the anticipated cohort of post-ICU patients that the Covid-19 pandemic will create over the next several months, across countries of the world. Patients and their caregivers need to be informed about what may be health consequences after the stay in the ICU as well as the long duration of limitations in functioning and the option to seek counseling and treatment.

The size of this cohort will be unprecedented as the disease surge will yield a large number of patients vulnerable to PICS and other less serious consequences of intensive care and mechanical ventilation. Many countries of the world are currently experiencing the catastrophic results of not preparing for a pandemic such as that we are experiencing now; it is urgent that countries do not fail to prepare for this major aftershock of the pandemic. Awareness of the inevitability of this aftershock is the first, and most important, step in the call for action. Professional societies across health care professions involved in rehabilitation and acute care can cooperate in the scaling up of the response within countries and regions. They may rely on tools and guidance provided by the World Health Organization (WHO) developed in response to "Rehabilitation 2030: A Call for Action".

It is vital to establish adequate screening opportunities (15). This may be done by a general practitioner alone or by a multiprofessional team consisting of a rehabilitation physician, a physical therapist, a psychologist, a critical care physician and others (16). The choice of a screening technique (including telemedicine and other e-health applications) depends on available resources, local healthcare infrastructure and availability of further rehabilitative interventions. There is strong evidence from systematic reviews that the availability and application of rehabilitation interventions are of primary importance to deal with the consequences of intensive care and mechanical ventilation (17–20). This has been supported by several randomized controlled trials as well (21–24).

It is therefore important to turn to the resources of rehabilitation hospitals and specialists in rehabilitation medicine and their multiprofessional teams who are equipped and experienced in providing the necessary interventions for mental, cognitive and physical impairments of the consequences of intensive care and mechanical ventilation. The capacity of both in- and outpatient rehabilitation of patients with PICS consequences will likely be sufficient, as public health social distancing measures might have eased their standard workload from traffic and work-related spinal cord injuries, head trauma and multi-trauma patients. Nevertheless, it remains important for policymakers to prepare to allocate additional resources to facilitate in- and outpatients' interventions for ICU survivors. Many rehabilitation institutions are already involved in acute care for patients with Covid-19. Some of them are equipped with ICU beds, and most of the rehabilitation institutions admit patients who have been discharged from ICU and are not yet ready to return home. Rehabilitation institutions may alleviate the enormous burden on acute hospitals and take care of a considerable number of patients.

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REFERENCES

- New COVID-19 Forecasts: US Hospitals Could Be Overwhelmed in the Second Week of April by Demand for ICU Beds, and US Deaths Could Total 81,000 by July [Internet] 2020 Mar 26 [cited 2020 Apr 8]. Available from: http:// www.healthdata.org/news-release/new-COVID-19-forecasts-us-hospitals-could-be-overwhelmed-second-weekapril-demand-icu.
- IHME COVID-19 health service utilization forecasting team. Forecasting COVID-19 impact on hospital bed-days, ICUdays, ventilator days and deaths by US state in the next 4 months. Medrxiv [Internet]. 2020 Mar 26 [cited 2020 Apr 8]. Available from: https://www.medrxiv.org/conten t/10.1101/2020.03.27.20043752v1.
- Xu B, Gutierrez B, Mekaru S, Sewalk K, Goodwin L, Loskill A, et al. Epidemiological data from the COVID-19 outbreak,

real-time case information. Sci Data 7 [Internet]. 2020 Mar 24 [cited 2020 Apr 8]; 106. Available from: https:// doi.org/10.1038/s41597-020-0448-0.

- Grasselli G, Pesenti A, Cecconi M. Critical Care Utilization for the COVID-19 Outbreak in Lombardy, Italy Early Experience and Forecast During an Emergency Response. JAMA 2020 Mar 13 [Epub ahead of print].
- Koh GC, Hoenig H. How should the rehabilitation community prepare for 2019-nCoV? Arch Phys Med Rehabil 2020 Mar 16 [Epub ahead of print].
- Edwards E. 'Post intensive-care syndrome': Why some COVID-19 patients may face problems even after recovery People who remain in the ICU for weeks may end up with memory problems and trouble thinking clearly. [Internet]. 2020 Mar [cited 2020 Apr 9]. Available from: https://www. nbcnews.com/health/health-news/post-intensive-caresyndrome-why-some-COVID-19-patients-may-n1166611.
- Flaatten H, Waldmann C.The Post-ICU syndrome, history and definition. In: Preiser J-C, Herridge M, Azoulay E, editors. Post-intensive care syndrome, lessons from the ICU. Cham, Switzerland: Springer; 2020, p. 3–12.
- Rawal G, Yadav S, Kumar R. Post -intensive care syndrome: an overview. J Transl Int Med 2017; 5: 90–92.
- 9. Harvey MA, Davidson JE. Post-intensive car syndrome: right care, right now...and later. Crit Care Med 2016; 44: 381–385.
- Zhou C, Wu L, Ni F, Ji W, Zhang, H. Critical illness polyneuropathy and myopathy: a systematic review. Neural Regen Res 2014; 9: 101–110.
- Herridge MS, Cheung AM, Tansey CM, Matte-Martyn A, Diaz-Granados N, Al-Saidi F, et al. One-year outcomes in survivors of the acute respiratory distress syndrome. N Engl J Med 2003; 348: 683–693.
- Needham DM, Davidson J, Cohen H, Hopkins RO, Weinert C, Wunsch H, et al. Improving long-term outcomes after discharge from intensive care unit: report from a stakeholders' conference. Crit Care Med 2012; 40: 502–509.
- Elliott D, Davidson JE, Harvey MA, Bemis-Dougherty A, Hopkins RO, Iwashyna TJ, et al. Exploring the scope of post-intensive care syndrome therapy and care: engagement of non-critical care providers and survivors in a second stakeholders meeting. Crit Care Med 2014; 42: 2518–2526.
- 14. Griffiths J, Hatch RA, Bishop J, Morgan K, Jenkinson C, Cuthbertson BH, et al. An exploration of social and economic outcome and associated health-related quality of life after critical illness in general intensive care unit survivors: a 21-month follow-up study. Crit Care 2013; 17: R100.
- Colbenson G A, Johnson A, Wilson ME. Post-intensive care syndrome: impact, prevention, and management. Breathe (Sheff) 2019; 15: 98–101.
- Van Der Schaaf M, Bakhshi-Raiez F, Van Der Steen M, Dongelmans DA, De Keizer NF. Recommendations for intensive care follow-up clinics: report from a survey and conference of Dutch intensive cares. Minerva Anestesiol 2015; 81: 135–144.
- 17. Stiller K. Physiotherapy in intensive care: an updated systematic review. Chest 2013; 144: 825–847.
- Kayambu G, Boots R, Paratz J. Physical therapy for the critically ill in the ICU: a systematic review and metaanalysis. Crit Care Med 2013; 41: 1543–1554.
- Mehlhorn J, Freytag A, Schmidt K, Brunkhorst FM, Graf J, Troitzsch U, et al. Rehabilitation interventions for postintensive care syndrome: a systematic review. Crit Care Med 2014; 42: 1263–1271.
- Castro-Avila AC, Serón P, Fan E, Gaete M, Mickan S. Effect of early rehabilitation during intensive care unit stay on functional status: Systematic review and meta-analysis. PLoS One 2015; 10: e0130722.
- 21. Greening NJ, Williams JE, Hussain SF, Harvey-Dunstan TC, Bankart MJ, Chaplin EJ, et al. An early rehabilitation

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intervention to enhance recovery during hospital admission for an exacerbation of chronic respiratory disease: randomised controlled trial. BMJ 2014; 349: g4315.

- Morris PE, Berry MJ, Files DC, Thompson JC, Hauser J, Flores L, et al. Standardized rehabilitation and hospital length of stay among patients with acute respiratory failure: a randomized clinical trial. JAMA 2016; 315: 2694–2702.
- 23. Moss M, Nordon-Craft A, Malone D, Van Pelt D, Frankel SK,

Warner ML, et al. A randomized trial of an intensive physical therapy program for patients with acute respiratory failure. Am J Respir Crit Care Med 2016; 193: 1101–1110.

 Schweickert WD, Pohlman MC, Pohlman AS, Nigos C, Pawlik AJ, Esbrook CL, et al. Early physical and occupational therapy in mechanically ventilated, critically ill patients: a randomised controlled trial. Lancet 2009; 373: 1874–1882.