



## **EDITORIALS**

## Covid-19 related hospital admissions in the United States: needs and outcomes

Lessons from rigorous inpatient studies will inform long term follow-up and continuing care

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Cohort studies and case series of patients with coronavirus disease 2019 (covid-19) have spread almost as quickly as the virus, <sup>1-5</sup> with inadequate methodological quality sometimes hampering thoughtful interpretation. For example, one New York cohort study initially reported a mortality of 88% for patients requiring mechanical ventilation before readers' responses led to a correction clarifying that the rate was 25% and would grow to an uncertain extent because 72% of patients were still in hospital.<sup>4</sup> In contrast, the two linked cohort studies from the United States provide rigorous outcome ascertainment and adjustment for case mix.<sup>67</sup> By delaying publication until most enrolled patients had confirmed outcomes and by using modern survival analysis methods these authors have greatly enhanced our understanding of the epidemiology of covid-19.

Lewnard and colleagues (doi:10.1136/bmj.m1923) report on 1840 people with covid-19 at Kaiser Permanente system hospitals in California and Washington state.<sup>6</sup> Median age was 61 years with 25% being 48 years or younger. The authors used survival analysis methods adjusted for patient characteristics to account for those who remained in hospital at the time of analysis. The study reported incidence of hospital admission (15-23 episodes per 100 000 cohort members in their insured populations), hospital length of stay (median 10.1 days), probability of intensive care unit (ICU) admission (41%), length of stay among ICU patients (median 10.5 days, and mortality. Mortality among all patients with completed hospital stays and outcomes was 18%, with risk factors including increased age and male sex.

The study also estimated dynamic changes in infection transmission rates by calculating the effective reproduction number  $(R_{\scriptscriptstyle E})$ . An  $R_{\scriptscriptstyle E}$  >greater than 1 indicates that an individual infects more than one other individual in a given population on a given day. The authors estimated a high  $R_{\scriptscriptstyle E}$  ranging from 1.31 to 2.49 for those acquiring infection on 1 March 2020 in the different regions covered by the study and lower  $R_{\scriptscriptstyle E}$  ranging

from 0.78 to 0.90 for those acquiring infection on 1 April 2020. This strong evidence of decreased transmission over time corresponds with enactment of social distancing policies.

Petrilli and colleagues (doi:10.1136/bmj.m1966) report on 5279 patients with covid-19 at an academic medical center in New York City. The cohort included patients testing positive for severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2), the virus responsible for covid-19, in any outpatient or emergency department setting. Median age of patients admitted to hospital was (62 years with 25% being 51 years or younger). The study outcomes included inpatient hospital stay (52%) and critical illness among those admitted to hospital (36%), representing a composite of ICU admission, mechanical ventilation, and hospital mortality or discharge to hospice care. Mortality among patients receiving mechanical ventilation was 60%. Of the 990 patients who developed critical illness, 399 (40%) were younger than 65 years with 91 (9%) patients being younger than 45 years. Of the entire cohort, outcome assessment was complete for 99.6% of patients, greatly enhancing confidence in these estimates.

Several patient characteristics in the New York study, including age, male sex, obesity, and heart failure were risks for more severe illness, mirroring earlier reports. The authors also provide compelling evidence confirming the predictive value of reduced oxygen saturation and increases in troponin, C reactive protein, and D-dimer levels near admission. As in previous reports, asthma and chronic obstructive pulmonary disease were notably absent from identified risk factors. Importantly, although emerging data show rates of covid-19 cases, hospital admissions, and deaths among African-American and Latino populations have been worse than among white populations, the authors did not find evidence of racial or ethnic disparities in critical illness or death once patients had been admitted to hospitals in their health system.

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Perhaps the most intriguing finding from the New York cohort was that risk of critical illness declined progressively over the study period, with a suggestion of declining mortality as well, without changes in risk of hospital admission. Several potential explanations worthy of future investigation include the influence of strain in hospital capacity on quality of care, allocation of resources, and disposition decisions in the emergency department<sup>10 11</sup>; changes in care delivery over time, such as proning in awake, non-intubated patients to avoid intubation<sup>12</sup> or better adherence to lung protective mechanical ventilation strategies; and changes in targeted therapy that might be beneficial (remdesivir<sup>13</sup> and anticoagulation<sup>14</sup>) or harmful (hydroxychloroquine<sup>15</sup>).

These studies provide the strongest evidence to date that for every covid-19 related hospital death in the US there have been four times as many survivors, typically requiring one to two weeks in hospital, and often in the ICU. The studies also provide rigorous confirmation of risk factors for poor outcomes that had been suggested in early reports. While older age is the greatest risk factor for adverse outcomes, including death, the burden of severe covid-19 illness is clearly borne across the lifespan, as most hospital admissions (53%<sup>7</sup> to 60%<sup>6</sup>) occurred in patients younger than 65 years. Early reports with incomplete follow-up are liable to considerable errors in reporting absolute event rates but often provide reasonable estimates of relative risks. These lessons will inform the transition to handling long term debility among hundreds of thousands of hospital survivors, deciding which patients can be managed safely at home, understanding risks for readmission, and developing effective strategies for long term care.

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