11-May-2020

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Factors associated with hospitalization and critical illness among 5,279 patients with COVID-19 disease in New York City: A prospective cohort study

Dear Dr. Horwitz (Leora)

Thank you for sending us your revision paper, which we sent back to one peer reviewer (who had no further concerns) and our statistical consultant (who had a few relatively minor points needing attention, as below). Please revise your manuscript in light of the remaining points, or let us know why you disagree. I look forward to hearing back from you as soon as possible.

Yours sincerely,

David Ludwig Professor David Ludwig Associate Research Editor The BMJ dludwig@bmj.com

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Reviewer: 1

This paper has been revised to take account of many of the statistical issues previously raised.

Follow-up has been extended and thus there are fewer patients with no set outcome data which reduces bias with respect to the critical illness logistic regression analysis. More details are provided about those who test negative. The analyses now include study week as a covariate and adjust for hospital. A new survival analysis (mortality) is also presented, and sensitivity analyses have also been carried out. The analysis involving critical illness in relation to all patients has been omitted, and the decision tree classification analysis has been removed from the paper.

There are just a few points that remain:

- 1. I agree that the very heterogenous group of tested subjects does not necessarily result in bias when comparing hospitalised vs non-hospitalised Covid-19 patients, although I am still concerned the decision to admit may be influenced by the 'reason/site for testing' which would, in turn, affect the hospitalised vs non-hospitalised comparison. It is useful to have the site data (Appendix Table S2) which shows that 59% of positive tests were carried out in the ED, and 96% of those hospitalised were tested in the ED. But it would be useful to know whether (in a sensitivity-type analysis) restricting the study sample to those tested in the ED gave similar results.
- 2. Tables 2-4. Because some of the significant multivariable-adjusted factors need to be interpreted carefully (eg a 'change' from a higher to a lower risk for hyperlipidema with statistical adjustment in Table 2) it would seem useful to add 'univariate' OR/HR results alongside the multivariable results here. This would highlight the sometimes complex relationship between a risk factor and hospitalisation/critical illness etc. Alternatively, specific factor relationships should be discussed more extensively in the text and the univariate results included in Supplementary Tables.

- 3. Out of interest, in the multivariable analyses, were interactions between any of the factors considered?
- 4. Table 1. I would suggest omitting the temperature and oxygen saturation results here as they are open to misinterpretation.
- 5. Figures. The time axis should include the units, ie. 'days'.

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Reviewer: 2

The authors have addressed my main comments and those of the other reviewers. The added figures showing cumulative incidence functions for mortality/discharge are of great value. Congratulations to the authors for a very informative study undertaken under extraordinarily difficult circumstances.

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