

**Judul : The Relationship D-dimer with severity and mortality in SARS-CoV-2 patients: A meta-analysis**

**Submission**

4/23/22, 6:32 PM

Email Airlangga University - Manuscript ID IJLH-06-20-0530



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**Manuscript ID IJLH-06-20-0530**

1 pesan

**Susmitha Raghubabu** <onbehalf@manuscriptcentral.com>

3 Juli 2020 11.47

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03-Jul-2020

Dear Dr Nugroho,

Your manuscript entitled "THE RELATIONSHIP OF D-DIMER WITH SEVERITY AND MORTALITY IN SARS-COV-2 PATIENTS: A META-ANALYSIS" has been successfully submitted online and is presently being given full consideration for publication in the International Journal of Laboratory Hematology.

Your manuscript ID is IJLH-06-20-0530.

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Co-authors: If you were unaware of this submission or do not agree with being listed as a co-author for this manuscript please contact the Editorial Office immediately.

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Best wishes,

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Primary Author: Johannes Nugroho

Manuscript: Relationship of D-dimer with severity and mortality in SARS-CoV-2 patients : A meta-analysis

Manuscript DOI: 10.1111/ijlh.13336

Manuscript ID: IJLH13336

Publication: *International Journal of Laboratory Hematology*

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4 Agustus 2020 20.13

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04-Aug-2020

Dear Dr Nugroho,

Manuscript ID IJLH-06-20-0530 entitled "THE RELATIONSHIP OF D-DIMER WITH SEVERITY AND MORTALITY IN SARS-COV-2 PATIENTS: A META-ANALYSIS", which you submitted to the International Journal of Laboratory Hematology, has now been reviewed. The comments of the reviewer(s) are included at the bottom of this letter.

Your manuscript has been recommended for publication subject to some major revisions suggested by the reviewer. Given the considerable pressure on space in the journal, it would also be helpful if you could delete any unnecessary material in order to reduce the length of your manuscript.

We recognize that the impact of the COVID-19 pandemic may affect your ability to return your revised manuscript to us within the requested time frame. If this is the case, please let us know

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Once again, thank you for submitting your manuscript to the International Journal of Laboratory Hematology and we look forward to receiving your revision.

Best wishes,

Susmitha Raghobabu  
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On behalf of the Editors

International Journal of Laboratory Hematology

Reviewer(s)' Comments to Author:

Reviewer: 1

Comments to the Author

I raise the following suggestions.

1. P4: Confirm that 'D-dimer' was not used as a search term. Does not negate the study, since likely to capture more papers, mostly unrelated to D-dimer, but should be clarified
2. Page 5 and elsewhere: "Only three studies had similar age of subjects in both groups." As D-dimer increases with age, and as COVID-19 affects more greatly those of older age, how did the authors control for the bias (higher D-dimer values expected in older people, which comprised more of the severe cases)?
3. Page 7 and elsewhere: "Pooled analysis by Shah et al. demonstrated that patient who had D-dimer value more than 0.5 mg/L had two fold higher risk of developing severe case and four-fold higher risk of mortality than those had D-dimer value less than 0.5 mg/L. Higher cut off value of D-dimer (>2 mg/L) was..." D-dimer has two main reporting units – fibrinogen equivalent units (FEU) and D-dimer units (DDU), which differ by a factor of ~2. What units were these reported values? Several errors in the literature have been identified in regards to D-dimer reporting units (Pubmed ID (PMID): 32432563). This should be mentioned in this report. Were such errors uncovered by the authors? What effect if any would such errors have on the authors' findings?
4. Table 1. Add another column next to "Median of D-dimer values" to identify if reported values are FEU, DDU or unknown/not reported in original paper.
5. Table 1: "Median of D-dimer values" seems to be sometimes with range; are all these really Medians, or sometimes Mean (i.e., those with +/- SD)?
6. Table 1: Add Table footer and define all abbreviations used in Table.

Minor

English can be improved slightly; examples: (a) page 2: "...using the PubMed..." (delete 'the'); "...were included in this meta-analysis. It revealed..." change to "...were included in this meta-analysis, which revealed..." (b) page 3: "then known caused by novel coronavirus named"; "...SARS-COV-2 infection. One of them was D-Dimer." (c) page 5: "None of the study". (d) page 7: "... (D-dimer and FDP) moderately or markedly elevated in all deaths." Has unclear meaning.

Reviewer: 2

Comments to the Author

This is a review of "THE RELATIONSHIP OF D-DIMER WITH SEVERITY AND MORTALITY IN SARS-COV-2 PATIENTS: A META-ANALYSIS" by Nugroho et al. It is a meta-analysis using PRISMA principles to evaluate D-dimer levels in severe vs non-severe disease, survivors vs non-survivors, and ICU vs non-ICU COVID-19 patients. I have the following specific comments:

1. The quality of the English is overall good but is below the quality for a scientific publication. However, there are occasional grammatical and spelling errors. For example, the "dimer" in D-dimer should be lower case throughout the text. The abbreviation for "ICU" is not defined on first use (abstract line 6). "Case" should be "cases" (abstract, results line 3), "patients with needs of ICU" is grammatically incorrect (abstract, results line 5). I don't think the word "case" is necessary in line 1 of the introduction. "Totally, we included only 29 studies ..." (results line 10). The entire manuscript should be reviewed for these and similar spelling and grammatical issues. Word or a similar word processing program may identify many of these errors.
2. The authors state that they used the PRISMA framework, which is the accepted standard for most academic journals. I don't see any obvious deviations from the PRISMA framework in the manuscript. However, I think it is good practice to include the PRISMA worksheet as a supplementary document with each item mapped to the specific page/line(s) of the manuscript. The worksheet is available online as a Word document.
3. Several studies in the manuscript were taken from a preprint source, which includes manuscripts undergoing peer review and non-peer review manuscripts voluntarily uploaded by researchers. I would recommend that the authors carefully review these studies to look for methodological flaws not detected by their quality analysis.
4. D-dimer may be the most-studied biomarker of disease severity in COVID-19, and a large number of studies have been published about it, as the authors correctly observe. Did this study produce any findings that add to our understanding of D-dimer in COVID-19?

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4 Agustus 2020 20.29

[Kutipan teks disembunyikan]

**Corrections successfully submitted for IJLH\_EV\_IJLH13336, Relationship of D-dimer with severity and mortality in SARS-CoV-2 patients : A meta-analysis.**

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8 September 2020 13.55

## Corrections successfully submitted

Dear **Johanes Nugroho**,

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Best regards,  
**Radhakrishnan, Deepan Thirumurthy**





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**Decision to accept revised manuscript ID IJLH-06-20-0530.R1**

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18 Agustus 2020 14.38

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18-Aug-2020

Dear Dr Nugroho,

We have pleasure in accepting your manuscript entitled "Relationship of D-dimer with severity and mortality in SARS-CoV-2 patients : A meta-analysis" in its current form for publication in the International Journal of Laboratory Hematology. The comments of the reviewer(s) who reviewed your manuscript are included at the foot of this letter.

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Best wishes,

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International Journal of Laboratory Hematology

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Reviewer(s)' Comments to Author:

Reviewer: 1

Comments to the Author  
no further suggestions

Reviewer: 2

Comments to the Author  
Thank you for your revision. My concerns have been addressed.

**nugroho eko putranto J.** <j.nugroho.eko@fk.unair.ac.id>

18 Agustus 2020 14.56

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1 September 2020 19.30

Dear Johannes Nugroho,

Article ID: IJLH13336  
Article DOI: 10.1111/ijlh.13336  
Internal Article ID: 16874223  
Article: Relationship of D-dimer with severity and mortality in SARS-CoV-2 patients : A meta-analysis  
Journal: International Journal of Laboratory Hematology

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6 Desember 2020 17.56

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15 September 2020 23.31

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