



Since January 2020 Elsevier has created a COVID-19 resource centre with free information in English and Mandarin on the novel coronavirus COVID-19. The COVID-19 resource centre is hosted on Elsevier Connect, the company's public news and information website.

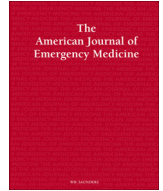
Elsevier hereby grants permission to make all its COVID-19-related research that is available on the COVID-19 resource centre - including this research content - immediately available in PubMed Central and other publicly funded repositories, such as the WHO COVID database with rights for unrestricted research re-use and analyses in any form or by any means with acknowledgement of the original source. These permissions are granted for free by Elsevier for as long as the COVID-19 resource centre remains active.



Contents lists available at ScienceDirect

American Journal of Emergency Medicine

journal homepage: www.elsevier.com/locate/ajem



Predictive performance of SOFA & qSOFA for in-hospital mortality in patients with severe novel coronavirus disease



Dear Editor,

We read the study by Liu et al. with great interest [1]. We believe that the authors' evaluation of the predictive performance of the Sequential Organ Failure Assessment (SOFA) and Quick Sequential Organ Failure Assessment (qSOFA) scores for hospital mortality in patients with severe and critical coronavirus 2019 (COVID-19) infection can be more easily applied at the bedside using likelihood ratios. Likelihood ratios are a measure of diagnostic accuracy that are derived from sensitivity and specificity. When patients test positive, the post-test probability that patients have the disease increases as the value of the positive likelihood ratio (LR+) becomes greater. For example, LR+ of 2, 5, and 10 increase post-test probability by roughly 15, 30%, and 45% respectively (though the increment in post-test probability decreases as it approaches 100%). Conversely, when patients test negative, the post-test probability that patients have the disease decreases as the value of the negative likelihood ratio (LR-) becomes smaller. LR- of 0.5, 0.2, and 0.1 decrease post-test probability by roughly 15, 30, and 45% respectively (though the increment in post-test probability decreases as it approaches 0) [2]. We calculated the LR+ and LR- of SOFA and qSOFA scores for hospital mortality in patients with severe and critical COVID-19 infection using the results from Liu et al.'s study (Table 1).

A SOFA score ≥ 3 is a moderately good positive predictor (LR+ 5.35, 95% confidence interval [CI] 3.43 to 8.36) and a strong negative predictor (LR- 0.12, 95% CI 0.03 to 0.45) for hospital mortality. Conversely, a qSOFA score ≥ 1 is a less positive predictor (LR+ 3.57, 95% CI 2.21 to 5.76) and a moderately good negative predictor (LR- 0.37, 95% CI 0.19 to 0.73). Using the pre-test probability for hospital mortality of (20/127) or 15.7% from the paper by Liu et al., the post-test probability of mortality in patients with SOFA ≥ 3 would be 49.9% and 2.2% for patients with a SOFA score 0 to 2, while the post-test probability for hospital mortality would be 39.3% in patients with qSOFA score ≥ 1 and 6.4% for patients with a qSOFA score of 0.

There are two key limitations that should be considered before applying these results in clinical practice. First, the likelihood ratios that we calculated may be overestimates as the data from Liu and colleagues were derived from an exploratory analysis of a single-center retrospective cohort using a data-driven approach [3]. A larger prospective cohort study to validate the optimal cutoff values reported in their study is warranted. Second, the strength of both the SOFA and qSOFA as

Table 1

Likelihood ratios of SOFA and qSOFA as predictors for in-hospital mortality of severe/critical COVID-19.

Models	Cutoff value	Sn (%)	Sp (%)	LR+ (95% CI)	LR- (95% CI)
All					
SOFA	1	100	50.47	2.02 (1.61, 2.41)	Not calculated
	2	95	71.96	3.39 (2.46, 4.66)	0.07 (0.01, 0.47)
	3	90	83.18	5.35	0.12 (0.03, 0.45)
	4	70	87.85	5.76 (3.21, 10)	0.34 (0.17, 0.67)
	5	55	94.39	9.8 (4.10, 23)	0.48 (0.29, 0.78)
	6	25	98.13	13 (2.79, 64)	0.76 (0.59, 0.99)
	7	20	98.13	11 (2.08, 53)	NS
	8	20	100	Not calculated	0.8 (0.63, 0.99)
qSOFA	1	70	80.37	3.57 (2.21, 5.76)	0.37 (0.19, 0.73)
	2	0	97.2	Not calculated	NS
	3	0	100	Not calculated	Not calculated
Age < 65 years					
SOFA	1	100	54.24	2.19 (1.72, 2.64)	Not calculated
	2	100	71.19	3.47 (2.49, 4.55)	Not calculated
	3	100	81.36	5.36 (3.47, 7.66)	Not calculated
	4	66.67	86.44	4.92 (2.78, 8.69)	0.39 (0.21, 0.72)
	5	33.33	93.22	4.92 (1.93, 13)	0.72 (0.52, 0.98)
	6	33.33	98.31	20 (4.09, 95)	0.68 (0.50, 0.93)
	7	33.33	98.31	20 (4.09, 95)	0.68 (0.50, 0.93)
	8	33.33	100	Not calculated	0.67 (0.49, 0.90)
qSOFA	1	66.67	76.27	2.81 (1.77, 4.45)	0.44 (0.23, 0.82)
	2	0	96.61	Not calculated	NS
	3	0	100	Not calculated	Not calculated
Age ≥ 65 years					
SOFA	1	100	45.83	1.85 (1.50, 2.17)	Not calculated
	2	94.12	72.92	3.48 (2.50, 4.83)	0.08 (0.01, 0.47)
	3	88.24	85.42	6.05 (3.72, 9.84)	0.14 (0.04, 0.46)
	4	70.59	89.58	6.77 (3.63, 13)	0.33 (0.17, 0.65)
	5	58.82	95.83	14 (5.30, 38)	0.43 (0.25, 0.73)
	6	23.53	97.92	11 (2.47, 52)	NS
	7	17.65	97.92	8.4 (1.69, 42)	NS
	8	17.65	100	Not calculated	NS
qSOFA	1	70.59	85.42	4.84 (2.82, 8.30)	0.34 (0.17, 0.68)
	2	0	97.92	Not calculated	NS
	3	0	100	Not calculated	Not calculated

Sn: Sensitivity, Sp: Specificity, LR+ : Positive Likelihood Ratio, LR-: Negative Likelihood Ratio, CI: Confidence Interval, NS: Not Significant.

Not Calculated: LR+ were not reported if there were no patients in one or more groups in the 2×2 table constructed to calculate the value because this would create an LR of 0 or infinity, which is unlikely.

predictors of mortality may be different today as new treatments have been shown to reduce mortality in patients with COVID-19 infection since this study was performed [4].

Declaration of Competing Interest

The authors declare no competing sources of interest.

References

- [1] Liu S, Yao N, Qiu Y, He C. Predictive performance of SOFA and qSOFA for in-hospital mortality in severe novel coronavirus disease. *Am J Emerg Med.* 2020;38(10): 2074–80. <https://doi.org/10.1016/j.ajem.2020.07.019>.
- [2] McGee S. Simplifying likelihood ratios. *J Gen Intern Med.* 2002;17(8):646–9. <https://doi.org/10.1046/j.1525-1497.2002.10750.x>.
- [3] Leeflang MM, Moons KG, Reitsma JB, Zwiderman AH. Bias in sensitivity and specificity caused by data-driven selection of optimal cutoff values: mechanisms, magnitude, and solutions. *Clin Chem.* 2008;54(4):729–37. <https://doi.org/10.1373/clinchem.2007.096032>.
- [4] Dennis JM, McGovern AP, Vollmer SJ, Mateen BA. Improving survival of critical care patients with coronavirus disease 2019 in England: a National Cohort Study, March to June 2020. *Crit Care Med.* 2021;49(2):209–14. <https://doi.org/10.1097/CCM.0000000000004747>.

Dominic Xiang Wang BHSc

*Schulich School of Medicine & Dentistry, Western University, London,
Ontario, Canada*

*Corresponding author at: Western University, 1151 Richmond St,
London, ON N6A 3K7, Canada.

E-mail address: dwang2021@meds.uwo.ca

Michael Ke Wang MD

*Department of Medicine, McMaster University, Hamilton, Ontario, Canada
Department of Health Research Methods, Evidence and Impact, McMaster
University, Hamilton, Ontario, Canada
Population Health Research Institute, McMaster University, Hamilton,
Ontario, Canada*

Bram Rochweg MD MSc

*Department of Medicine, McMaster University, Hamilton, Ontario, Canada
Department of Health Research Methods, Evidence and Impact, McMaster
University, Hamilton, Ontario, Canada*

Karen E.A. Burns MD

*Interdepartmental Division of Critical Care Medicine, University of Toronto,
Toronto, Ontario, Canada
Li Ka Shing Knowledge Institute, Unity Health Toronto - St. Michael's
Hospital, Toronto, Ontario, Canada*

25 February 2021