Manuscript Title	Coronavirus Disease-2019 (COVID-19): An overview and a clinical update
Authors	Arunkumar Krishnan, James P. Hamilton, Saleh A. Alqahtani, Tinsay A.Woreta
Journal	World Journal of Clinical Cases
Manuscript Number	59037
Comments Received	18 Oct 2020
Responses Sent	22 Oct 2020

S. No.	Reviewer / Section	Comments	Response
Peer Review Comments			
1	Reviewer # 1 / Scientific quality	Section/subsection headings and hierarchy are confusing. Please clarify and/or reorganize sections and subsections.	All headings have now been aligned properly with the introduction of 'heading styles' (Heading 1 and Heading 2).
		Many discussion points are binary and dichotomous (eg, immune vs. non-immune; up vs. down-regulation; old vs. young; etc.). It is a bit misleading because many biological characteristics form or represent a continuum. There are many environmental, dietary, and lifestyle factors that influence the microbiome (in both gut lumen and tissue), immune system, pathogenic mechanisms, and response to vaccination. The authors should discuss these points; influence of those factors, eg, smoking, alcohol, diet, obesity, microbiome, etc. on immunity. Personal differences in immune response likely result from these factors as well as our genetic differences. Currently, there is lack of data in those areas. In these contexts, as a future	Suggested points have been added with adequate references in the section discussing vaccines.

S. No.	Reviewer / Section	Comments	Response
		direction, research on dietary / lifestyle factors, microbiome, immunity, and personalized molecular biomarkers is needed. The authors should discuss molecular pathological epidemiology (MPE), which can investigate those factors in relation to molecular pathologies, immunity, and clinical outcomes. MPE, its strengths and challenges have been discussed in J Pathol 2019, Annu Rev Pathol 2019, etc. MPE research can be a promising direction and improve prediction of response to vaccine or other forms of immune-based intervention.	
Editorial Office Comments			
2	Science Editor / Scientific quality / Academic norms and rules	The highest single-source similarity index showed to be 7% in the CrossCheck report. The authors need to rephrase the repeated parts.	The texts that showed to have 7% similarity with published sources have now been rephrased.
3	Figures	The authors did not provide original pictures. Please provide the original figure documents. Please prepare and arrange the figures using PowerPoint to ensure that all graphs or arrows or text portions can be reprocessed by the editor.	Figures are moved to a PPT and submitted with the prescribed file name.
		Please provide decomposable Figures (whose parts are all movable and editable), organize them into a single PowerPoint file, and submit as "**59037-Figures.ppt**" on the system. The figures should be uploaded to the file destination of "Image File".	
4	Tables	Please provide decomposable Tables (whose parts are all movable and editable), organize them into a single Word file, and submit as "**59037-Tables.docx**" on the system. The tables should be	Tables are separately provided in a single Word document with the specified file name.

S. No.	Reviewer / Section	Comments	Response
		uploaded to the file destination of "Table File".	
5	Copyright	Please click and download the Copyright License Agreement Form. Subsequently, a PDF (scanned) version of the Copyright License Agreement Form that has been signed by all authors should be uploaded to the file destination of 'Copyright License Agreement'.	Copyright form has been duly signed and submitted.
6	Disclosure	Please click and download the fillable ICMJE Form for Disclosure of Potential Conflicts of Interest (PDF), and fill it in. The Corresponding Author is responsible for filling out this form. Once filled out completely, the Conflict-of-Interest Disclosure Form should be uploaded to the file destination of 'Conflict-of-Interest Disclosure Form'.	ICMJE COI Disclosure Form has been duly filled and submitted.
7	References	Please revise the references according to the Format for references guidelines, and be sure to edit the reference using the reference auto-analyser.	References have been updated per the references guidelines.

Answering reviewers for second round reviewc

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Comments Received	03 Nov 2020
Responses Sent	13 Nov 2020

S. No.	Reviewer / Section Comments	Response
1.	This is a generally well-written review on COVID-19. The topic is of very high interest. This paper can be improved in following points. Section/subsection headings and hierarchy are confusing. Please clarify and/or reorganize sections and subsections.	Thank you for the comment and valuable suggestion. We have arranged the section and subsection with the numbers to avoid confusion.
2.	Many discussion points are binary and dichotomous (eg, immune vs. non-immune; up vs. down-regulation; old vs. young; etc.). It is a bit misleading because many biological characteristics form or represent a continuum.	Thank you for the comment. We understand your concern. We have explained the current status based on the available literature. COVID-19 is a rapidly evolving pandemic as we are still learning something new every day.
3.	There are many environmental, dietary, and lifestyle factors that influence the microbiome (in both gut lumen and tissue), immune system, pathogenic mechanisms, and response to vaccination. Currently, there is lack of data in those areas. In these contexts, as a future direction, research on dietary / lifestyle factors, microbiome, immunity, and personalized molecular biomarkers is needed. The authors should discuss molecular pathological epidemiology (MPE), which can investigate those factors in relation to molecular pathologies, immunity, and clinical outcomes. MPE, its strengths and challenges have been discussed in J Pathol 2019, Annu Rev Pathol 2019, etc.	Thank you for your comments and valuable suggestions. We have explained the section with molecular pathological epidemiology. We have included a statement about the microbiome in the vaccination section.

	MPE research can be a promising direction and improve prediction of response to vaccine or other forms of immune-based intervention	
	The authors should discuss these points; influence of those factors, eg, smoking, alcohol, diet, obesity, microbiome, etc. on immunity. Personal differences in immune response likely result from these factors as well as our genetic differences.	Thank you for your comments and suggestions. We have included a section to explain the role of lifestyle interventions, where we explained the details of the role of nutrition, the effect of Microbiota on COVID-19, the role of Probiotics, and the effect of smoking and alcohol. We have explained about the influence of obesity.