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PEER-REVIEW REPORT

Name of journal: World Journal of Cardiology

Manuscript NO: 65348

Title: Red Blood Cell Distribution Width in Elderly Hospitalized Patients with

Cardiovascular Disease

Reviewer's code: 05824934 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Singapore

Author's Country/Territory: Greece

Manuscript submission date: 2021-03-06

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-03-07 23:10

Reviewer performed review: 2021-03-11 00:36

Review time: 3 Days and 1 Hour

Scientific quality	[] Grade A: Excellent [Y] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
Language quality	[Y] Grade A: Priority publishing [] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [] Accept (General priority) [Y] Minor revision [] Major revision [] Rejection
Re-review	[Y]Yes []No
Peer-reviewer statements	Peer-Review: [] Anonymous [Y] Onymous Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

The paper investigated the red blood cell distribution width (RDW) in elderly patients with cardiovascular disease, and the results show that RDW was significantly elevated in these patients. The findings in this paper are interesting and worth to be published. Yet, the discussion part on immunity and inflammation has room for improvement. As what was pointed out by the authors, hypertension, coronary artery disease, diabetes mellitus and chronic obstructive pulmonary disease are among the most common morbidities in patients with elevated RDW. These morbidities all belong to metabolic syndromes. So, instead of considering poor nutritional status as mentioned in the manuscript, one may suspect the contribution of overnutrition to chronic diseases or Human immunity is an essential part of the integrated nutrition acquisition and supply mechanism of the human body [1], especially when the person falls ill and inflammation is initiated in certain tissues [2, 3]. The human immune system initiates inflammation in response to tissue damage [4]. Inflammation is a protective immunological reaction to remove the injurious stimuli, and remove the damaged tissue as well as initiate the tissue healing process [4]. Yet, overnutrition will prevent the tissue healing process from happening. This is because the nutrition from the breakdown of the damaged tissue by inflammation together with the excessive nutrition already inside the body will be mostly turned into lipid intermediates, causes lipotoxicity (the deposition of lipid intermediates in non-adipose tissue, leading to cellular dysfunction and death) [5] in healthy non-adipose tissues, and induces further tissue damage. The breakdown of non-adipose tissues and formation of lipid intermediates results in a vicious cycle. Thus, the overnutrition situation is worsened by the loss of lean body mass, coupled with perpetual chronic inflammation. The following reference may be included in the manuscript to provide a more broad picture of the topic discussed in this manuscript:



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References: [1] Yu, LG (2021) Restoring Good Health in Elderly with Diverse Gut Microbiome and Food Intake Restriction to Combat COVID-19. Indian Journal of Microbiology, 61, 104–107. https://doi.org/10.1007/s12088-020-00913-3 [2] van Niekerk G, Isaacs AW, Nell T, Engelbrecht AM (2016) Sickness-Associated Anorexia: Mother Nature's Idea of Immunonutrition? Mediators of Inflammation 2016 Article Number: 8071539. DOI: 10.1155/2016/8071539. [3] Arabi YM, Reintam BA, Preiser JC (2019) Less is more in nutrition: critically ill patients are starving but not hungry. Intensive Care Medicine, 45(11), pp. 1629-1631. DOI: 10.1007/s00134-019-05765-0 [4] Costantini S, Sharma A and Colonna G (2011). The Value of the Cytokinome Profile, Inflammatory Diseases - A Modern Perspective, Dr. Amit Nagal (Ed.), ISBN: 978-953-307-444-3, InTech, Available

http://www.intechopen.com/books/inflammatory-diseases-a-modern-perspective/the -value-of-the-cytokinome-profile [5] Garbarino J, Sturley SL (2009) Saturated with fat: new perspectives on lipotoxicity. Curr Opin Clin Nutr Metab Care 12:110–116. DOI: 10.1097/mco.0b013e32832182ee



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PEER-REVIEW REPORT

Name of journal: World Journal of Cardiology

Manuscript NO: 65348

Title: Red Blood Cell Distribution Width in Elderly Hospitalized Patients with

Cardiovascular Disease

Reviewer's code: 02446694 **Position:** Editorial Board

Academic degree: FACC, FAHA, MD, PhD

Professional title: Doctor

Reviewer's Country/Territory: Japan

Author's Country/Territory: Greece

Manuscript submission date: 2021-03-06

Reviewer chosen by: Man Liu

Reviewer accepted review: 2021-03-12 19:28

Reviewer performed review: 2021-03-20 23:07

Review time: 8 Days and 3 Hours

Scientific quality	[] Grade A: Excellent [] Grade B: Very good [] Grade C: Good [Y] Grade D: Fair [] Grade E: Do not publish
Language quality	[] Grade A: Priority publishing [Y] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
Conclusion	[] Accept (High priority) [] Accept (General priority) [] Minor revision [Y] Major revision [] Rejection
Re-review	[Y]Yes []No
Peer-reviewer	Peer-Review: [Y] Anonymous [] Onymous
statements	Conflicts-of-Interest: [] Yes [Y] No



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SPECIFIC COMMENTS TO AUTHORS

This study seems to be interesting, however, there are one important problem and several comments. #1 This study was performed prospectively and this study needs the informed consent from patients and the ethics committee certificate of study protocol. #2 The authors should show the reasons of selecting patients of age > 65 years as the elderly patients. #3 In the second paragraph of "Discussion" section, beginning with "In the current study RDW---", the meaning of this paragraph was unclear. #4 Some values were expressed with the second decimal place, however, these expressions were meaningless to some extent. The authors should express values with proper decimal place. #5 The whole of manuscript seems to be too lengthy. The authors should shorten the whole of manuscript, especially in the "Discussion" section.



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RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Cardiology

Manuscript NO: 65348

Title: Red Blood Cell Distribution Width in Elderly Hospitalized Patients with

Cardiovascular Disease

Reviewer's code: 05824934 Position: Peer Reviewer Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Singapore

Author's Country/Territory: Greece

Manuscript submission date: 2021-03-06

Reviewer chosen by: Man Liu

Reviewer accepted review: 2021-04-13 07:04

Reviewer performed review: 2021-04-14 01:06

Review time: 18 Hours

[] Grade A: Excellent [Y] Grade B: Very good [] Grade C: Good [] Grade D: Fair [] Grade E: Do not publish
[Y] Grade A: Priority publishing [] Grade B: Minor language polishing [] Grade C: A great deal of language polishing [] Grade D: Rejection
[] Accept (High priority) [] Accept (General priority) [Y] Minor revision [] Major revision [] Rejection
Peer-Review: [] Anonymous [Y] Onymous Conflicts-of-Interest: [] Yes [Y] No



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As the authors have addressed all my concerns in my previous review report, I have no further comments except the following minor revision suggestions: 1. Page 10, last line, "poor nutritional status" may be replaced by "nutritional imbalance"; 2. Page 11, line 1, "or nutrient deficiencies (eg, iron, vitamin B12, or folate deficiency) that are associated with anisocytosis [14]." may be replaced by "expressed by micronutrient deficiencies (eg, iron, vitamin B12, or folate deficiency) that are associated with anisocytosis [14], and excess of macronutrients." 3. Page 11, line 5, "Overnutrition" may be replaced by "Macronutrient surplus"