Answers to the Reviewers

investigation.

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2 Reviewer #1: 3 **Scientific Quality:** Grade C (Good) 4 **Language Quality:** Grade B (Minor language polishing) 5 **Conclusion:** Minor revision 6 7 **Specific Comments to Authors:** This study mainly explored the importance of vitamin D supplementation in patients with autoimmune hepatitis. Through the relationship 8 between AIH and the immune system, as well as the relationship between vitamin D 9 10 and the immune system, this article leads to the correlation between the immune regulation, anti-inflammatory, antioxidant, and anti-fiber effects of vitamin D and the 11 prevention and treatment of AIH. However, there is a lot of irrelevant content. 12 Moveover, I need to highlight several important considerations regarding this research. 13 First of all, according to clinical manifestations, autoimmune liver diseases are divided 14 into three types. Different types of autoimmune liver diseases have different clinical 15 manifestations and pathological changes in the liver. Will the prognostic value of 16 17 vitamin D supplementation vary depending on these subtypes? At the same time, vitamin D deficiency has been described in various liver diseases and non-liver diseases 18 19 and other healthy people, so what is the reference vitamin D supplement dosage for 20 patients with autoimmune liver disease, and how to give appropriate vitamin D 21 supplementation for patients with different severity of liver disease? 22 Thank you for your comments and suggestions. We deleted some not relevant 23 sentences. We focused on the effect of vitamin D on autoimmune hepatitis retrieving 24 useful data from a recent systematic review. However, a number of questions remain 25 not answered because as I indicated in my title. It is a topic that needs further 26

28 29 Reviewer #2: 30 **Scientific Quality:** Grade C (Good) 31 **Language Quality:** Grade A (Priority publishing) 32 **Conclusion:** Minor revision 33 **Specific Comments to Authors:** The article is quite interesting and open a new research 34 domain. The conclusion did not reflect the content of the article. Several conclusion are 35 written but the substrate is not presented in the article (e.g. Vitamin D improves 36 glucocorticoids, increases the production of glutathione, and inhibits hepatic stellate 37 cells. Vitamin D may influence the histological severity of AIH, advanced liver fibrosis, 38 and the need for liver transplantation). I suggest to include these results in order to 39 40 have these conclusion. The author should also mention what is the recommendation for supplementation of vitamin D in autoimmune hepatitis. In this way the author could 41 correlate better the tile of the article with the main content. 42 43 Thank you for your comments and suggestions. According to your suggestions, we 44 specifically focused on the effect of vitamin D on autoimmune hepatitis retrieving 45 46 useful data from a recent systematic review. The recommendation for vitamin D for 47 patients with autoimmune hepatitis remains elusive. I retrived papers on the effect of 48 vitamin D on single basic functions of the hepatocytes. A number of questions remain not answered because as I indicated in my title. It is a topic that needs further 49 investigation. I changed the title as well. 50 51 52

53 Reviewer #3: 54 **Scientific Quality:** Grade E (Do not publish) 55 **Language Quality:** Grade C (A great deal of language polishing) 56 **Conclusion:** Rejection 57 58 **Specific Comments to Authors:** The title of this letter is 'Vitamin D is a critical 59 supplement in patients suffering from autoimmune hepatitis'. The author argues that 60 vitamin plays an important role in the immune system. A few important issues need to 61 be addressed. 1. Title: The title is vague. Does vitamin D play an important role in the pathogenesis of autoimmune hepatitis? What is being 'critical'? 2. The author did not 62 present any convincing data linking the pathogenesis of autoimmune hepatitis with 63 vitamin D deficiency. The mere presence of vitamin D receptor on the surface of 64 peripheral blood mononuclear cells does not automatically make it important in 65 regulating the immune system. No reference is cited to support this claim. 3. In the last 66 paragraph, the author claims vitamin D has immunoregulatory, anti-inflammatory, 67 anti-oxidative, and anti-fibrotic effects, affecting the outcome of immune-mediated 68 diseases. There was no supporting references, including specific reference linking 69 70 vitamin D and autoimmune hepatitis. 4. Similarly, the statement linking vitamin D may influence the histological severity of AIH, advanced liver fibrosis and the need of liver 71 transplantation is also not supported by any appropriate references. 5. Finally, the 72 73 usage of English language needs to be improved. 74 Thank you for your comments and suggestions. We changed the title and retrieve single 75 papers on the effect of hepatocytes and specifically on NAFLD and an important article 76 on Turkish patients affected with autoimmune hepatitis and treated with vitamin D 77 supplementation. We listed more references suggesting how vitamin D can act on the 78 79 body influencing glutathione and glucocorticoid response. Since this is a letter to the

editor, we found that an expansion of the text would make the manuscript a review. We

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added a microphotograph of a patient with autoimmune hepatitis with remarkable interface hepatitis. We improved the English language (grammar and spellings) and apologize for previous version. Reviewer #4: Scientific Quality: Grade B (Very good) Language Quality: Grade B (Minor language polishing) **Conclusion:** Accept (General priority) Specific Comments to Authors: This is a well written manuscript. Focus more on immunomodulatory function of vitamin D rather than other effects e.g. skeletal. Thank you for your comments and suggestions. We focused on autoimmune hepatitis, added a microphotograph and expanded the references suggesting the potential support of vitamin D supplementation for patients suffereing from autoimmune hepatitis.