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**Name of journal:** World Journal of Gastroenterology

**ESPS manuscript NO:** 30332

**Title:** To screen or not to screen? Celiac antibodies in liver diseases

November 14<sup>th</sup>, 2016

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: WJG - Narciso-Schiavon - Full manuscript - Celiac Liver - reviewed).

*Response from the authors:*

*We would like to thank you for providing invaluable advices regarding our manuscript. We hope we will be able to address your comments properly. The manuscript has been improved according to the suggestions of reviewers. See specific answers to reviewers bellow.*

**Name of journal:** World Journal of Gastroenterology

**ESPS manuscript NO:** 30332

**Title:** To screen or not to screen? Celiac antibodies in liver diseases

**Reviewer's code:** 03646970

**Reviewer's country:** United States

**Science editor:** Jing Yu

**Date sent for review:** 2016-09-28 14:35

**Date reviewed:** 2016-10-19 13:04

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> [ Y] Accept
<input type="checkbox"/> [ Y] Grade B: Very good	<input type="checkbox"/> [ Y] Grade B: Minor language	<input type="checkbox"/> [ ] The same title	<input type="checkbox"/> [ ] High priority for
<input type="checkbox"/> [ ] Grade C: Good	polishing	<input type="checkbox"/> [ ] Duplicate publication	publication
<input type="checkbox"/> [ ] Grade D: Fair	<input type="checkbox"/> [ ] Grade C: A great deal of	<input type="checkbox"/> [ ] Plagiarism	<input type="checkbox"/> [ ] Rejection
<input type="checkbox"/> [ ] Grade E: Poor	language polishing	<input type="checkbox"/> [ ] No	<input type="checkbox"/> [ ] Minor revision
	<input type="checkbox"/> [ ] Grade D: Rejected	BPG Search:	<input type="checkbox"/> [ ] Major revision
		<input type="checkbox"/> [ ] The same title	
		<input type="checkbox"/> [ ] Duplicate publication	
		<input type="checkbox"/> [ ] Plagiarism	
		<input type="checkbox"/> [ ] No	

## COMMENTS TO AUTHORS

Well written manuscript addressing the needed variety of disorders. Minor grammar or spelling changes are suggested in the manuscript.

*R: Grammar and spelling have been professionally reviewed by Enago™, Crimson Interactive Inc.*

## ESPS PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**ESPS manuscript NO:** 30332

**Title:** To screen or not to screen? Celiac antibodies in liver diseases

**Reviewer's code:** 00068093

**Reviewer's country:** Turkey

**Science editor:** Jing Yu

**Date sent for review:** 2016-09-28 14:35

**Date reviewed:** 2016-11-02 00:04

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input checked="" type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	Google Search:	<input checked="" type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

## COMMENTS TO AUTHORS

No

*R: Thank you.*

## ESPS PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**ESPS manuscript NO:** 30332

**Title:** To screen or not to screen? Celiac antibodies in liver diseases

**Reviewer's code:** 00003692

**Reviewer's country:** Canada

**Science editor:** Jing Yu

**Date sent for review:** 2016-09-28 14:35

**Date reviewed:** 2016-11-02 00:49

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

## COMMENTS TO AUTHORS

Very nice review. Dr. Hugh Freeman

*R: Thank you.*

## ESPS PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**ESPS manuscript NO:** 30332

**Title:** To screen or not to screen? Celiac antibodies in liver diseases

**Reviewer's code:** 03699905

**Reviewer's country:** Germany

**Science editor:** Jing Yu

**Date sent for review:** 2016-09-28 14:35

**Date reviewed:** 2016-11-04 20:16

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good		<input type="checkbox"/> Duplicate publication	
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade E: Poor		<input type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
	<input type="checkbox"/> Grade D: Rejected	BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input type="checkbox"/> No	

### COMMENTS TO AUTHORS

Congratulation for the authors for the review.

#### MAJOR COMMENTS;

1. Page 2, bottom, unexplained/idiopathic transaminasemia is an indication to rule out CD.

*R: What we meant was: "but is often overlooked in the differential diagnosis of liver diseases". We have modified the text accordingly.*

2. Page 3, 2nd paragraph; AGA are not autoimmune antibodies.

*R: Thank you. We have modified the text.*

3. Page 3 middle: Combination test are better than single test for CD diagnosis. *Lerner A. Serological Diagnosis of Celiac Disease –Moving Beyond the Tip of the Iceberg. International Journal of Celiac Disease. Editorial. 2014;2:64-66.*

*R: We have modified the text and added the above-mentioned reference.*

4. Page 3 middle:For false+/- IgA tTg please site:*A Lerner, P Jeremias, T Matthias. Outside of Normal Limits:False Positive/Negative Anti TG2 Autoantibodies. Internat J Celiac Disease, 2015;3:87-90.*

*R: We have added the above-mentioned reference to the text.*

5. Page 3 middle-lower:There are serological kits that combine IgA+IgG anti tTg or neo-epitope tTg :*A Lerner, P Jeremias, S Neidhofer, T Matthias. Antibodies against neo-epitope tTg complexed to gliadin are different and more reliable then anti-tTg for the diagnosis of pediatric celiac disease. J Immunol Methods. 2016;429:15-20.*

*R: We have added the information to the text and added the above-mentioned reference.*

6. Page 3 bottom:Neo-epitope tTg should be mentioned as serological marker of CD:*Lerner A, Neidhofer S, Matthias T. Serological markers and/or intestinal biopsies in the case-finding of celiac disease. Editorial, Internat. J Celiac dis. 2015;3:53-55. Lerner A. More novel diagnostic antibodies for celiac disease. Expert Rev of Gastroenterol & Hepatol 2016 May 30;1-2. [Epub ahead of print] PubMed PMID: 27237317.*

*R: We have added the information to the text and added the above-mentioned reference.*

7. page 5 Material & methods:The reviewers screened the literature of which years, or for how many decades? Is only PubMed was evaluated? Are only English manuscripts were considered? Why "transaminasemia" was not screened? Were abstracts considered? (see ref 40)

*R: Thank you for your comments. We are sorry for any inconvenience. We searched for MeSH terms and transaminasemia was not screened. Abstracts were considered when we consulted references of selected articles. We have corrected the text.*

8. Page 9 upper part:Will site for false+/- tTg: *A Lerner, P Jeremias, T Matthias. Outside of Normal Limits: False Positive/Negative Anti TG2 Autoantibodies. Internat J Celiac Disease, 2015;3:87-90.*

*R: We have added the above-mentioned reference to the text.*

9. Page 10 upper part; common antigenic basis. please consider using molecular mimicry

*R: We have modified the text, as suggested.*

10. Page c10 bottom:Other organs tTg:*Lerner A, Neidhofer S, Matthias T. Transglutaminase 2 and Anti Transglutaminase 2 Autoantibodies in Celiac Disease and Beyond: TG2 Double-Edged Sword: Gut and Extraintestinal Involvement. Immunome Research, 2015;11:101-105.*

*R: We have added the above-mentioned reference to the text.*

11. Page 10, bottom .Please site references of GFD preventing autoimmune diseases.

*R: We have added appropriate references to the text.*

12. page 17, bottom:"pretreatment normal liver enzyme" not clear enough.

*R: We have modified the text: "Interestingly, in those patients in whom pretreatment liver enzyme levels are normal, we do observe a significant decrease in their serum levels with a gluten-free diet."*

13. Page 18 ,HCV. please mention the strong association between HCV and autoimmune diseases in general

*R: HCV might be involved in the breaking of tolerance to self-antigens and therefore in triggering auto reactivity. HCV has been implicated both in the triggering of autoimmune diseases and in the development of autoantibodies.*

14. Page 21, last paragraph should be In summary, to summarize the main take home messages of the review.

*R: We modified the abstract, as suggested.*

#### **MINOR REMARKS**

1. English should be improve

*R: Grammar and spelling have been professionally reviewed by Enago™, Crimson Interactive Inc.*

2. Abbreviations should be used consistently, where nemtioned (CD,AIH etc).

*R: We have modified the text and abbreviations accordingly.*