

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 04913316

Position: Peer Reviewer

Academic degree: MD, PhD

Professional title: Full Professor

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: AI Technique

Reviewer accepted review: 2020-03-28 06:55

Reviewer performed review: 2020-03-29 08:52

Review time: 1 Day and 1 Hour

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 Grade E: Do not publish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



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SPECIFIC COMMENTS TO AUTHORS

This is an interesting review which explores thoroughly the extra gastric manifestations of *Helicobacter pylori* infection. The review is well written' references are adequate and the manuscript is overall informative. I only have a few suggestions which might help improve the manuscript. Comments 1) The Authors correctly quoted a recent review dealing with extra gastric manifestations written by Gravina et al. However this reference is quoted twice (as reference 7 and as reference 51). One of these reference should be eliminated. 2) In the paragraph dealing with ophthalmological manifestations I suggest to discuss and quote in the reference list the study by Cotticelli L et al (Eur J Ophthalmol 2006; 16:264-278) showing an association between *H. pylori* infection and central serous chorioretinopathy. 3) In the Introduction, when talking about the route of transmission of the infection, I suggest to quote the manuscript by Sgambato D et al recently published in United European Gastroenterol J 2018; 6:1470-1476.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 02954663

Position: Peer Reviewer

Academic degree: PhD

Professional title: Chief Doctor

Reviewer's Country/Territory: Hungary

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: Ruo-Yu Ma (Quit in 2020)

Reviewer accepted review: 2020-04-07 06:08

Reviewer performed review: 2020-04-08 08:04

Review time: 1 Day and 1 Hour

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



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SPECIFIC COMMENTS TO AUTHORS

Ironically, enough, after discovery of *Helicobacter pylori*, most of diseases appeared to be associated with this infection and this created confusion. I am afraid that the manuscript submitted, unfortunately, will not clear this confusion. In most of cases, it is true that some statistical degree of association between prevalence of *H. pylori* and the given disease was demonstrated, but this, nevertheless, does not mean a cause-effect relationship and this fact is not at all emphasized in the manuscript. Although the manuscript is intended to present the extragastric manifestations of the *H. pylori* infection, some conditions such as inflammatory bowel diseases, GERD/esophageal carcinoma are also included and discussed, while in all these the role of bacteria could be even protective, but this is not discussed. Vitamin B12 deficiency is a consequence of gastric mucosal atrophy caused by the infection and autoimmune reaction, so that it is not an extragastric manifestation. Pediatric issues are omitted. Association between *H. pylori* and colon polyps/CRC, chronic pancreatitis/pancreatic cancer is also forgotten. Linguistic polishing is also required. In figure 2, "intestinal bowel disease" is probably irritable bowel disease or IBD? The position statements regarding eradication of *H. pylori* in extragastric diseases as published in several recent guidelines (Maastricht/Florence V, Toronto, AGA, Brazilian, Chinese, Mexican, Bangkok) is not commented.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 00503417

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor, Professor

Reviewer's Country/Territory: India

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: Ruo-Yu Ma (Quit in 2020)

Reviewer accepted review: 2020-04-08 07:52

Reviewer performed review: 2020-04-08 08:57

Review time: 1 Hour

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

SPECIFIC COMMENTS TO AUTHORS

This is a good review of the subject. However, there are recent reviews on the same topic published elsewhere (Franceschi F, Covino M, Roubaud Baudron C. Review: *Helicobacter pylori* and extragastric diseases. *Helicobacter*. 2019 Sep;24 Suppl 1:e12636) as well as in this journal itself (Gravina AG, Zagari RM, De Musis C, Romano L, Loguercio C, Romano M. *Helicobacter pylori* and extragastric diseases: A review. *World J Gastroenterol*. 2018 Aug 7; 24(29): 3204–3221). The authors should state how the current review adds to the recent ones. Also, they can add value to their review if they weigh the evidence for and against individual associations and rate the associations based on the evidence.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 03010350

Position: Peer Reviewer

Academic degree: MD, PhD

Professional title: Professor

Reviewer's Country/Territory: Russia

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: AI Technique

Reviewer accepted review: 2020-03-29 07:09

Reviewer performed review: 2020-04-12 16:50

Review time: 14 Days and 9 Hours

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

SPECIFIC COMMENTS TO AUTHORS

I read this article for quite some time. The article is very interesting, but in many ways debatable, as well as all pathophysiological reviews. Nevertheless, the authors presented updated material on the problem. The citation of modern meta-analyses is impressive. I recommend publishing this article.

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 00051367

Position: Editorial Board

Academic degree: MD

Professional title: Doctor, Professor

Reviewer's Country/Territory: Italy

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: Ruo-Yu Ma (Quit in 2020)

Reviewer accepted review: 2020-04-14 13:33

Reviewer performed review: 2020-04-14 17:54

Review time: 4 Hours

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

SPECIFIC COMMENTS TO AUTHORS

It is a interesting and sufficient review. It may be interesting for readers if the authors could briefly comment the condition halitosis. It think that the papers should include a table summarizing evidences of each condition (scarce sufficient, relevant ...).

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 55671

Title: Helicobacter pylori infection: beyond gastric manifestations

Reviewer's code: 01799104

Position: Editorial Board

Academic degree: AGAF, MD

Professional title: Associate Professor, Doctor

Reviewer's Country/Territory: Taiwan

Author's Country/Territory: Brazil

Manuscript submission date: 2020-03-28

Reviewer chosen by: Ruo-Yu Ma (Quit in 2020)

Reviewer accepted review: 2020-04-08 08:28

Reviewer performed review: 2020-04-19 10:03

Review time: 11 Days and 1 Hour

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



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SPECIFIC COMMENTS TO AUTHORS

The authors did an explicit review of all the related diseases. Most of the disease are positively related to and few are inversely related to Hp infection. Maybe I would suggest that you put a summary in your conclusion or table to further state which is either causative or protective. Furthermore, several diseases have been put into treatment guideline, such as idiopathic thrombocytopenia or iron deficiency anemia and the authors may emphasize these points. Failure to thrive for children has been reported but the fact is still controversial. You may add another session because there are sufficient studies in the literature though you have mentioned under the session of iron deficiency anemia. There are some queer symbols in the sessions of NAFLD and Parkinson's disease.