



PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45797

Title: Personalized medicine in functional gastrointestinal disorders: Understanding pathogenesis to increase diagnostic and treatment efficacy

Reviewer’s code: 02989927

Reviewer’s country: Brazil

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-18

Date reviewed: 2019-01-18

Review time: 4 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good		<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	(General priority)	Peer-reviewer’s expertise on the topic of the manuscript:
<input type="checkbox"/> Grade E: Do not publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Minor revision	<input type="checkbox"/> Advanced
		<input checked="" type="checkbox"/> Major revision	<input checked="" type="checkbox"/> General
		<input type="checkbox"/> Rejection	<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

45797 Personalized Medicine in Functional Gastrointestinal Disorders: Time to Tailor the Treatment on the Basis of the Pathogenesis? by Wang & Camilleri, 2019. This is an interesting narrative review on the phenotypes of functional gastrointestinal disorders



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and the therapeutic application of its drug-related pharmacogenomics. After reviewing main types of functional gastrointestinal disorders, the authors report therapeutic advances of functional gastrointestinal disorders in pharmacogenomics and drug pharmacokinetics, mutations and genetic variants, mucosal gene expression of functional gastrointestinal disorders. They conclude that patient-based research can provide additional personalization of diagnosis and management for patients with gastrointestinal disorders, with the aid of better understanding of the pathophysiology and clinically-applicable diagnostic tests. The title is provocative, but does not reflect the global content of the article at all. In general, the type of review should be stated in the title. The text is organized into a section of Background, Description of some functional gastrointestinal disorders (in terms of putative mechanism and the therapeutics), and indications of Precision Medicine. I would suggest a Table to summarize each type of gastrointestinal phenotype, along with a Table with recent advances of therapeutic alternatives. The authors can underscore the level of evidence for each treatment option. Because there is no critical assessment of the quality of existing evidence, I would replace the section of Conclusion by "Comments". A brief description of the limitations of the present review is welcome as well. Hope these recommendations can improve the text editing of the article.

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45797

Title: Personalized medicine in functional gastrointestinal disorders: Understanding pathogenesis to increase diagnostic and treatment efficacy

Reviewer's code: 03000421

Reviewer's country: France

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-18

Date reviewed: 2019-01-19

Review time: 4 Hours, 1 Day

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input checked="" type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good		<input checked="" type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	(General priority)	Peer-reviewer's expertise on the topic of the manuscript:
<input type="checkbox"/> Grade E: Do not publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Minor revision	<input checked="" type="checkbox"/> Advanced
		<input type="checkbox"/> Major revision	<input type="checkbox"/> General
		<input type="checkbox"/> Rejection	<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

This is an interesting review (see comments in the attached file)

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45797

Title: Personalized medicine in functional gastrointestinal disorders: Understanding pathogenesis to increase diagnostic and treatment efficacy

Reviewer's code: 00504353

Reviewer's country: Japan

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-18

Date reviewed: 2019-01-21

Review time: 23 Hours, 2 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language	(High priority)	<input type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
<input type="checkbox"/> Grade E: Do not	language polishing	<input type="checkbox"/> Minor revision	topic of the manuscript:
publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input type="checkbox"/> Advanced
		<input type="checkbox"/> Rejection	<input type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

This review looks written very well.

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45797

Title: Personalized medicine in functional gastrointestinal disorders: Understanding pathogenesis to increase diagnostic and treatment efficacy

Reviewer's code: 03317125

Reviewer's country: China

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-18

Date reviewed: 2019-01-27

Review time: 3 Hours, 9 Days

Table with 4 columns: SCIENTIFIC QUALITY, LANGUAGE QUALITY, CONCLUSION, PEER-REVIEWER STATEMENTS. It contains checkboxes for various quality grades (A-E), conclusion types (Accept, Revision, Rejection), and reviewer statements regarding expertise and conflicts of interest.

SPECIFIC COMMENTS TO AUTHORS

In this review article Wang et al, has put together the recent developments in specific mechanisms and personalized treatment of functional gastrointestinal disorders. The article is informative and provides a nice overview. However, this reviewer has a few



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suggestions that the authors should consider addressing. 1. Recent information suggests that the association of functional gastrointestinal disorders and psychological impairment may be more fundamental than was previously believed. The rationale for psychological therapy in the treatment of functional gastrointestinal disorders should be also emphasized to give readers a wider view of the problem. 2. Functional gastrointestinal disorders are divided into functional upper gastrointestinal symptoms and functional lower gastrointestinal symptoms in the article, and related mechanisms have been listed. However, most of them are related to one or a type of symptom without mentioning possible mechanisms and treatments for multiple symptoms. 3. The authors mention that nutritional factors, including FODMAPs are associated with predominant pain or bloating, but miss out on dietary fibers. A couple of sentences regarding its association with functional gastrointestinal disorders should be included in the review. 4. This review proposes an individualized approach to the management of functional gastrointestinal disorders on the basis of the pathogenesis. In fact, a table listing all the specific mechanisms and their personalized treatment of functional gastrointestinal disorders would be really informative. 5. Accumulating studies have revealed the associations between gut dysbiosis and functional gastrointestinal disorders, and manipulating the microbial populations with therapeutic intent has become a hot topic of functional gastrointestinal disorders research. I would add some information about them.

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PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45797

Title: Personalized medicine in functional gastrointestinal disorders: Understanding pathogenesis to increase diagnostic and treatment efficacy

Reviewer's code: 01557050

Reviewer's country: Japan

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-18

Date reviewed: 2019-01-29

Review time: 20 Hours, 10 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	(High priority)	<input type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejection	(General priority)	Peer-reviewer's expertise on the topic of the manuscript:
<input type="checkbox"/> Grade E: Do not publish		<input type="checkbox"/> Minor revision	<input type="checkbox"/> Advanced
		<input type="checkbox"/> Major revision	<input type="checkbox"/> General
		<input type="checkbox"/> Rejection	<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
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SPECIFIC COMMENTS TO AUTHORS

Dr. Wang and Dr. Camilleri reviewed 'Personalized Medicine in Functional Gastrointestinal Disorders: Time to Tailor the Treatment on the Basis of the Pathogenesis?'

The manuscript is informative and well-presented. The reviewer has no comments.



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