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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 25990

Title: The regulation of serotonin transporter in irritable bowel syndrome

Reviewer's code: 02156774

Reviewer's country: Taiwan

Science editor: Ya-Juan Ma

Date sent for review: 2016-03-28 17:33

Date reviewed: 2016-04-11 13:45

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"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

#25990, The regulation of serotonin transporter in irritable bowel syndrome Dr. Jin et al. conducted a review on the role of serotonin transporter (SERT) on irritable bowel syndrome (IBS). IBS could be a disorder that may harm the quality of daily life. Considering the role of serotonin on GI system, it is clear that the role of serotonergic system is important. I have some concerns listed below:

1. It should be noticed that IBS could be related with psychosomatic effect, and other comorbids with psychiatry disorder, such as depression and anxiety. These disorders are also related with altered serotonin system. This issue should be introduced.
2. I appreciated the authors introduced some treatment strategy in the section gut microbiota. It has been reviewed that antidepressant, especial the SSRI could be effective treatments for IBS, also. This issue should be introduced in an expended section for the potential treatments.



ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 25990

Title: The regulation of serotonin transporter in irritable bowel syndrome

Reviewer’s code: 01427097

Reviewer’s country: Japan

Science editor: Ya-Juan Ma

Date sent for review: 2016-03-28 17:33

Date reviewed: 2016-04-15 22:40

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 checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input checked="" 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"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

Authors reviewed IBS focusing on SERT by quoting many papers. However the association of 5-HT or SERT with IBS seems to be different based on subtypes of IBS. Because SERT plays an important role regulating 5-HT signaling which generally induce to diarrhea, authors should introduce the papers indicating the specific subtypes. e.g. All subjects of ref 35 were post-infectious IBS and IBS-D except one subjects with IBS-D. The subjects of Ref 91 investigating miRNA-24 were IBS-D. It is difficult to distinguish the difference between the IBS-C and IBS-D in the some parts. In the introduction of the manuscript, it is better to describe that the other important factors except 5-HT signaling also play major important roles in immunity and inflammation and Gut microbiota of the IBS patients. Moreover authors should describe why authors focused on SERT. Ref 76 written in Chinese is better to be deleted.

ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

ESPS manuscript NO: 25990

Title: The regulation of serotonin transporter in irritable bowel syndrome

Reviewer's code: 02531171

Reviewer's country: Ireland

Science editor: Ya-Juan Ma

Date sent for review: 2016-03-28 17:33

Date reviewed: 2016-04-26 19:21

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 checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input checked="" 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"/> Grade D: Rejected	<input checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
		BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

COMMENTS TO AUTHORS

The review entitled 'The regulation of serotonin transporter in irritable bowel syndrome' by Du-chen Jin and colleagues describes the role of the serotonin transporter in the pathophysiology of irritable bowel syndrome. This is a well written, comprehensive review of SERT in functional bowel disorders. The standard of written English is good but there are a number of grammatical and typographical and citation errors throughout the text which need to be corrected. Introduction: when describing the subtypes of IBS, post-infective IBS should also be mentioned (as it is discussed later in the review). When describing the mechanisms underlying IBS, 'visceral hypersensitivity [15,20], intestinal barrier dysfunction [21] and gut motility disorder [15,16,22]' are really more characteristic symptoms of the disorder, rather than mechanisms. Pg 10: explain what Shugan decoction is and its proposed mechanism of action (and how this relates to SERT gene expression). Pg 12: some factual errors relating to the microbiome eg. 'variety of creative attempts, especially probiotics, prebiotics, antibiotics and FMT, to increase the relative abundance of probiotics (like Lactobacilli and Bifidobacteria, etc.)' change probiotics to 'commensals' Pg 12: '...found that probiotics Faecalibacterium prausnitzii strain A2-165 or its supernatant, which was also a type of commensal



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bacterium..’ The supernatant would not be a bacterium, rather the secretory product of that bacterium. The final paragraph relating to antibiotics and FMT and SERT in IBS needs to be strengthened (or removed). Pg 13: ‘SERT expression, such as EGF [127], basic fibroblast growth factor (bFGF) [128], nerve growth factor (NGF) [129], etc.’ Complete the sentence – don’t use etc. Pg 14: The final two sections on the role of growth factors and SERT in IBS and the role of other factors, need to be strengthened. The data presented doesn’t currently support a role for these factors in IBS. There is no mention of current or previous IBS treatments which targeted the serotonergic system and how they could be improved upon or side effects avoided in the future. Relating to therapeutic strategies, the importance of each of the subtypes in relation to serotonin should be made more clear.