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## PEER-REVIEW REPORT

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

**Manuscript NO:** 36829

**Title:** Regulation of the intestinal immune system by flavonoids and its utility in chronic inflammatory bowel disease

**Reviewer's code:** 02741591

**Reviewer's country:** Egypt

**Science editor:** Ke Chen

**Date sent for review:** 2017-10-26

**Date reviewed:** 2017-10-26

**Review time:** 2 Hours

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

Abstract: The abstract is truncated (141 words). It could better be expanded. The word 'dysbalance' should be (imbalance). English language needs polishing and revision by a native speaker. Ah receptor COMMENT: Fully expand the abbreviation at its first appearance in the abstract (aryl hydrocarbon) receptor. Introduction: The first 20 lines (2 paragraphs) should include references. The structure should be improved so that an introductory paragraph is presented before the subsequent subheadings. The review lacks in-depth of the presented literature. The references are overused for lengthy paragraphs. The subheading 'Flavonoids regulate the Ah-receptors', includes 12 lines cited by one reference. The literature presented in the last paragraph of the review does not comply with the preceding subheading and does not confer evidence of inhibition of intestinal inflammation by flavonoids. The references are few



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for a review article. As mentioned at the end of the conclusion, an original article clinical study design would be more appealing than a shortly presented incompletely informative review.



## PEER-REVIEW REPORT

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

**Manuscript NO:** 36829

**Title:** Regulation of the intestinal immune system by flavonoids and its utility in chronic inflammatory bowel disease

**Reviewer's code:** 02822560

**Reviewer's country:** Japan

**Science editor:** Ke Chen

**Date sent for review:** 2017-10-26

**Date reviewed:** 2017-11-02

**Review time:** 6 Days

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 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 checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		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

The main cause of IBDs is not fully understood, but an immune system disturbance as well as genetic disposition and environmental factors are associated with their pathogenesis. Significant antioxidant and free radical scavenging properties of flavonoids are well known and additionally, accumulating evidence in animal models or cell cultures suggest that flavonoids can inhibit inflammatory pathway via nuclear transcription factor, including NF- $\kappa$ B, ameliorating the intensity of inflammation in the digestive tract. In this mini-review, the authors briefly explain the Ah-receptor regulated by flavonoids as their main inhibitory effects against intestinal inflammation, but it seems difficult for the readers of this journal to understand the following two items: cytochrome, including CYP1A1, and PERM system, at each section of "FLAVONOIDS AND MICROSOMAL ENZYMES" and "FLAVONOIDS AND PERM", because they had



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written them too concisely. Their abnormality in patients with IBD and therapeutic benefits from some flavonoids should be explained in parallel.



**PEER-REVIEW REPORT**

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

**Manuscript NO:** 36829

**Title:** Regulation of the intestinal immune system by flavonoids and its utility in chronic inflammatory bowel disease

**Reviewer's code:** 02998194

**Reviewer's country:** Greece

**Science editor:** Ke Chen

**Date sent for review:** 2017-10-26

**Date reviewed:** 2017-11-08

**Review time:** 12 Days

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"/> 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 checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		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**

Dear Authors This is a nice mini review but still needs amendment For instance, in introductory part there are solid statements with no references. Furthermore the CRC part seems irrelevant In Anti-inflammatory effects of flavonoids part, the sentence Some flavonoids such as tea flavonoids (apigenin and epigallocatechin gallate) have been found to prevent the activation of immune cells and its effectors (chemokines, TNF- and cytokines) has no reference In Flavonoids and microsomal enzymes part, the PERM system needs explanation In Flavonoids and PERM, the conclusion is not clear and from the above Evidence of inhibition of intestinal inflammation by flavonoids In the sentence There are multiple experimental studies which indicate the utility of flavonoids in models of chronic inflammatory bowel disease[12-13]. These two references are not multiple