



PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Oncology

Manuscript NO: 51370

Title: Caffeine and Its Main Targets of Colorectal Cancer

Reviewer's code: 00077674

Position: Associate Editor

Academic degree: DVM, PhD

Professional title: Academic Research, Professor

Reviewer's country: United States

Author's country: China

Reviewer chosen by: Artificial Intelligence Technique

Reviewer accepted review: 2019-09-13 13:09

Reviewer performed review: 2019-09-13 13:49

Review time: 1 Hour

| 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 is an excellent review that summarizes the potential effects of caffeine and the



**Baishideng
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7041 Koll Center Parkway, Suite
160, Pleasanton, CA 94566, USA
Telephone: +1-925-223-8242
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involved mechanisms on colo-rectal cancer. However, the authors have overlooked an important aspect of caffeine: Due to its ability to increase intracellular cAMP, caffeine enhances the downstream effects of beta-adrenergic receptors that are coupled to the stimulatory G-protein Gs which increases cAMP by activating adenylyl cyclase. There are numerous reports in the literature that beta-adrenergic receptor signaling in response to their physiological agonists epinephrine and norepinephrine or in response to psychological stress that increases the systemic levels of these stress neurotransmitters stimulate the growth and angiogenesis of colo-rectal cancer. These aspects and the stimulatory effects of caffeine on these pathways need to be added to the review.

INITIAL REVIEW OF THE MANUSCRIPT

Google Search:

- The same title
- Duplicate publication
- Plagiarism
- No

BPG Search:

- The same title
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- Plagiarism
- No



PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Oncology

Manuscript NO: 51370

Title: Caffeine and Its Main Targets of Colorectal Cancer

Reviewer's code: 02942856

Position: Peer Reviewer

Academic degree: BM BCh

Professional title: Associate Professor, Attending Doctor

Reviewer's country: Taiwan

Author's country: China

Reviewer chosen by: Artificial Intelligence Technique

Reviewer accepted review: 2019-09-13 06:06

Reviewer performed review: 2019-09-15 03:59

Review time: 1 Day and 21 Hours

| SCIENTIFIC QUALITY | LANGUAGE QUALITY | CONCLUSION | PEER-REVIEWER STATEMENTS |
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| <input checked="" type="checkbox"/> Grade C: Good | polishing | <input type="checkbox"/> Accept | <input type="checkbox"/> Onymous |
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| publish | <input type="checkbox"/> Grade D: Rejection | <input type="checkbox"/> Major revision | <input type="checkbox"/> Advanced |
| | | <input type="checkbox"/> Rejection | <input checked="" type="checkbox"/> General |
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| | | | Conflicts-of-Interest: |
| | | | <input type="checkbox"/> Yes |
| | | | <input checked="" type="checkbox"/> No |

SPECIFIC COMMENTS TO AUTHORS

WJG 51370 Caffeine and Its Main Targets of Colorectal Cancer In this review article,



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Cui et al. review the molecular mechanisms implicated in the influences of caffeine on colorectal cancer and intestinal homeostasis, including the modulation of gut microbiota. This review has encompassed the various aspects of the interactions between caffeine and the colon and presents nicely with delicate figures. There are, however, some issues to be clarified: Major comments 1. Fig.3 shows that caffeine only blocks the action of PDE and reduces the synthesis of 5'-AMP without influencing the main pathways (cAMP AKt, ERK, p38MAPK.....) that promotes the carcinogenesis of colorectal cancer cells. 2. Likewise, the presentation of Fig.4 may mislead the readers to feel that caffeine is "promoting" the progression of colorectal cancer cells via the substances shown in this figure. 3. Please adjust the Fig.5, because it may also mislead the readers to feel that caffeine positively regulates TGF- β , EGFR.....then promotes the growth of colorectal cancer cells. Minor comments 1. Please delete ".....according to this article" at the end of figure legend 5, because this is a review article based upon the previous literature, but not an original article with novel findings.

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BPG Search:

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[Y] No



PEER-REVIEW REPORT

Name of journal: World Journal of Gastrointestinal Oncology

Manuscript NO: 51370

Title: Caffeine and Its Main Targets of Colorectal Cancer

Reviewer's code: 02573214

Position: Editorial Board

Academic degree: MD

Professional title: Full Professor, Surgeon

Reviewer's country: Italy

Author's country: China

Reviewer chosen by: Artificial Intelligence Technique

Reviewer accepted review: 2019-09-12 17:19

Reviewer performed review: 2019-09-18 14:40

Review time: 5 Days and 21 Hours

| SCIENTIFIC QUALITY | LANGUAGE QUALITY | CONCLUSION | PEER-REVIEWER STATEMENTS |
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| | | | <input type="checkbox"/> No expertise |
| | | | Conflicts-of-Interest: |
| | | | <input type="checkbox"/> Yes |
| | | | <input checked="" type="checkbox"/> No |

SPECIFIC COMMENTS TO AUTHORS

This manuscript is a good description of the state of art about the Caffeine and Its Main



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Targets of Colorectal Cancer.

INITIAL REVIEW OF THE MANUSCRIPT

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