



**PEER-REVIEW REPORT**

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

**Manuscript NO:** 46843

**Title:** Role of epigenetics in the transformation of inflammation into colorectal cancer

**Reviewer's code:** 02631746

**Reviewer's country:** United States

**Science editor:** Jia-Ping Yan

**Reviewer accepted review:** 2019-03-02 18:29

**Reviewer performed review:** 2019-03-03 18:49

**Review time:** 1 Day

| 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 | (High priority)                                    | <input checked="" type="checkbox"/> Anonymous |
| <input checked="" 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 checked="" type="checkbox"/> Minor revision | topic of the manuscript:                      |
| publish   | <input type="checkbox"/> Grade D: Rejection                 | <input type="checkbox"/> Major revision            | <input checked="" type="checkbox"/> Advanced  |
|   |   | <input type="checkbox"/> Rejection                 | <input type="checkbox"/> General              |
|   |   |  | <input type="checkbox"/> No expertise         |
|   |   |  | Conflicts-of-Interest:                        |
|   |   |  | <input type="checkbox"/> Yes                  |
|   |   |  | <input checked="" type="checkbox"/> No        |

**SPECIFIC COMMENTS TO AUTHORS**

Interesting review. It would have been better had the authors discussed relationship between genetic and epigenetic events in the pathobiology of inflammation induced colon cancer.



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**PEER-REVIEW REPORT**

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

**Manuscript NO:** 46843

**Title:** Role of epigenetics in the transformation of inflammation into colorectal cancer

**Reviewer's code:** 02575643

**Reviewer's country:** Italy

**Science editor:** Jia-Ping Yan

**Reviewer accepted review:** 2019-04-04 11:02

**Reviewer performed review:** 2019-04-04 11:27

**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 checked="" type="checkbox"/> Grade B: Minor language | (High priority)                                    | <input type="checkbox"/> Anonymous          |
| <input checked="" type="checkbox"/> Grade C: Good | polishing   | <input type="checkbox"/> Accept                    | <input checked="" 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 checked="" 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 checked="" type="checkbox"/> General |
|   |   |  | <input type="checkbox"/> No expertise       |
|   |   |  | Conflicts-of-Interest:                      |
|   |   |  | <input type="checkbox"/> Yes                |
|   |   |  | <input checked="" type="checkbox"/> No      |

**SPECIFIC COMMENTS TO AUTHORS**

Interetsing review. Many suggestions 1-This review needs tables for each subheading, and figures to explain schematically the action of the different etiology mechanisms conceptually leading from inflammation to cancer development. 2-The action of chronic administration of anti inflammatory drugs in reducing the prevalence of colorectal



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cancer should be reported in a separate sub heading., adding a meta analysis which shows the results of the several prospective studies which have demonstrated the reduced prevalence of colorectal cancer in patients taking anti inflammatory therapy. 3-The role of inflammation in colorectal cancer developmet opens many new therapeutic horizons, including the role of new anti inflammatory drugs. The Authors should add a sub-heading on this possibility.In this context, the Authors should analyze the possible side-effects, including the possibility of increased mortality and morbidity in head trauma, which national statistics have shown increased in the last years in the lederly (probably related to the action of anti aggregant-anti inflammatory chronic administration in the elderly). 4-I would freccomend to give special attention to the relationship of mucionous adenocarcinoma and inflammation. In patients with inflammatory bowel disease, there is a higher prevalenece of mucinous secreting adenocarcinoma, which is not evident in sporadic and familiar cases. 5-Colorectal cancer etiology implies many genetic factors. Inflammation seems to play a major role. 6-Among the many actions analyzed in the review, the Authors should include the imcreased production of inflammatory cytokines and growth factors which are present in the inflammatory cascade. The sub heading should also include the specific role of NK-f and the canonical and not canonical activation of this factor.

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**PEER-REVIEW REPORT**

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

**Manuscript NO:** 46843

**Title:** Role of epigenetics in the transformation of inflammation into colorectal cancer

**Reviewer's code:** 02495872

**Reviewer's country:** United States

**Science editor:** Jia-Ping Yan

**Reviewer accepted review:** 2019-04-04 13:40

**Reviewer performed review:** 2019-04-04 16:02

**Review time:** 2 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 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**

The idea and concept of this review is excellent and clinically important. Cancer and inflammatory diseases have been classified as non-communicable diseases for decades. Both diseases have characteristics of immune reactions, which are principally identical, but differing in important aspects. Still, the idea of connections between cancer and



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inflammation is often overlooked. The authors adequately introduced the situation and subsequently methodically went through individual epigenetic factors. All were substantially discussed. The paper is well-referenced and I am sure that it will serve as an excellent source of information on this subject.

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## PEER-REVIEW REPORT

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

**Manuscript NO:** 46843

**Title:** Role of epigenetics in the transformation of inflammation into colorectal cancer

**Reviewer's code:** 03471268

**Reviewer's country:** Japan

**Science editor:** Jia-Ping Yan

**Reviewer accepted review:** 2019-04-06 08:14

**Reviewer performed review:** 2019-04-12 02:49

**Review time:** 5 Days and 18 Hours

| 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 type="checkbox"/> Grade B: Very good       | <input type="checkbox"/> Grade B: Minor language                 | (High priority)                                    | <input checked="" type="checkbox"/> Anonymous |
| <input checked="" 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 checked="" 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 checked="" type="checkbox"/> General   |
|   |  |  | <input type="checkbox"/> No expertise         |
|   |  |  | Conflicts-of-Interest:                        |
|   |  |  | <input type="checkbox"/> Yes                  |
|   |  |  | <input checked="" type="checkbox"/> No        |

### SPECIFIC COMMENTS TO AUTHORS

This review is well written and summarized for the chronic inflammation-induced epigenetic changing-mediated CRC. If the authors could add the solution as well the future perspective for it, it will be more informative and helpful for readers.



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