

Format for ANSWERING REVIEWERS



June 15, 2015

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 19304-review.doc).

Title: Colon cancer and the epidermal growth factor receptor: Current treatment paradigms, the importance of diet, and the role of chemoprevention

Author: Baldeep S Pabla, Marc Bissonnette, Vani J Konda

Name of Journal: *World Journal of Clinical Oncology*

ESPS Manuscript NO: 19304

The manuscript has been improved according to the suggestions of reviewers:

Reviewer 02534290:

- Comments:
 - "Very good review on the subject, many recent aspects of the problem exposed and an integrative view given at the end."
- Response:
 - Thank you for your review of our paper. We appreciate that you noticed the integrative approach we took at looking at the role of the EGFR as it relates to current treatment and preventative modalities, with an eye towards the future.

Reviewer 00467700:

- Comments:
 - "Very good review, every aspect of egfr about colon cancer is well exposed."
- Response:
 - Thank you for your review of our paper.

Reviewer 02546401:

- Comments:
 - "The manuscript reviewed risk factors, chemoprevention and current treatment modalities for colorectal cancer, focusing on EGFR pathway. The study concept is clear. And the manuscript is well and concisely written. However, several major points need to point out. 1. English editing is required. 2. The title of the manuscript should be revised to make it clear and highlighted. 3. There is no abstract in the manuscript. 4. The authors should add the newest literatures in the manuscript."
- Response:
 - Thank you for your review of our paper. We have added a running title and the abstract we submitted is now clearly designated in our final submission. We

believe our references to be up to date and inclusive of all the current literature relevant to this topic.

Reviewer 02445433

- Comments:
 - "It will be helpful divide the review into sections and subsections (not all included EGFR, Diet and not Chemoprevention). In particular, the relation between EGFR and diet should contain: 1.the role of western diet; 2.the role of calcium (that is missing); 3. the role of phytochemicals. In the "approach to management" section, the gender difference in the prognostic significance of KRAS and BRAF mutations (Wangefjord S et al, 2013) must be included. In the " risk factors"section, the role of the stochastic stem cell division on the incidence of CRC would be helpful to cite and discuss (Tomassetti and Vogelstein, 2015). The role of miRNAand EGFR, targeting and CRC prognosis, is missing and will be valuable to introduce. It must be added a conclusion section where the main evidences on the interventions of prevention of CRC, in relation to the review content, will be resumed and may be discussed."
- Response:
 - Thank you for your review of our paper. We agree that the role of western diet and phytochemicals as it relates to the EGFR is important, and we highlighted the biochemical pathways implicated in western diets high in fat and the role of ginseng, green tea, and curcumin accordingly. We decided to keep our focus narrow and did not discuss the role of calcium or the role of stochastic stem cell division. We have included a brief description of the role of miRNAs in EGFR expression in accordance with your suggestions. We agree that the prognostic significance of KRAS and BRAF mutations as they relate to gender is an interesting area of research and have included some of these findings in our revised manuscript as well.