

**World J Gastrointest Oncol**

**Manuscript #: 55397**

**Title: Espresso coffee, Caffeine and Colon Cancer**

To the editor:

We revise references according to instruction. We provide PMID and DOI for each ref.

Reviewer #1:

*In this letter, the authors made a summary about the published article “Caffeine and its main targets of colorectal cancer.” by Cui and coworkers and pointed out that caffeine-containing drinks were difficult to standardize epidemiological studies in humans thus cannot draw consistent conclusions. Therefore, they would like to make additional contributions to the previous study that polyphenols might play a role on the association between espresso coffee and the development colon cancer via antioxidant effect. What’s more, there’s a difference between men and women in carcinogenesis mediated by oxidative stress. This supplement to the previous study is of great significance and it provides ideas for further studies. And the manuscript organized well.*

We thanks reviewer for her/his nice comment.

Reviewer #2:

*This letter to the editor showed a similar view to the original paper with some additional thoughtful comments.*

We thanks reviewer for her/his comment.

Reviewer #3:

*This letter provides a good extension of the cited review. The authers choose Espresso coffee to further investigate the relationship of caffeine’s effects on CRC and the effect of antioxidants in coffee which may be different between men and women. Due to antioxidant activity of coffee has a close relationship with its concentration, it would be better to discuss the*

*concentration of coffee or antioxidants that will be used in prospective randomized study. Finally, I hope Italy can overcome the epidemic as soon as possible. FORZA ITALIA!*

First of all we would like to thank reviewer for her/his support in this very difficult moment for our Country. We really appreciate.

Regarding suggestion about “*to discuss the concentration of coffee or antioxidants that will be used in prospective randomized study*” we add a comment on the on-going study (page 4 line 6-16 of the revised manuscript). We also add a comment on *prospective randomized study*. on page 5 line 1-3 of the revised manuscript.