

Title: Mechanisms of resistance to anti-EGFR inhibitors in metastatic Colorectal Cancer.

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Name of Journal: World Journal of Gastroenterology

Dear Editor,

Please find our revised manuscript entitled "Mechanisms of resistance to anti-EGFR inhibitors in metastatic Colorectal Cancer" according to the reviewer's comments.

Responses to the reviewer's comments;

- 1) The reviewer 00722674 suggested correcting some orthographic mistakes, to provide the running title, the key words and to re-edit figures. He/she also suggested reducing the words of the title.
- 1) We have made the required changes.

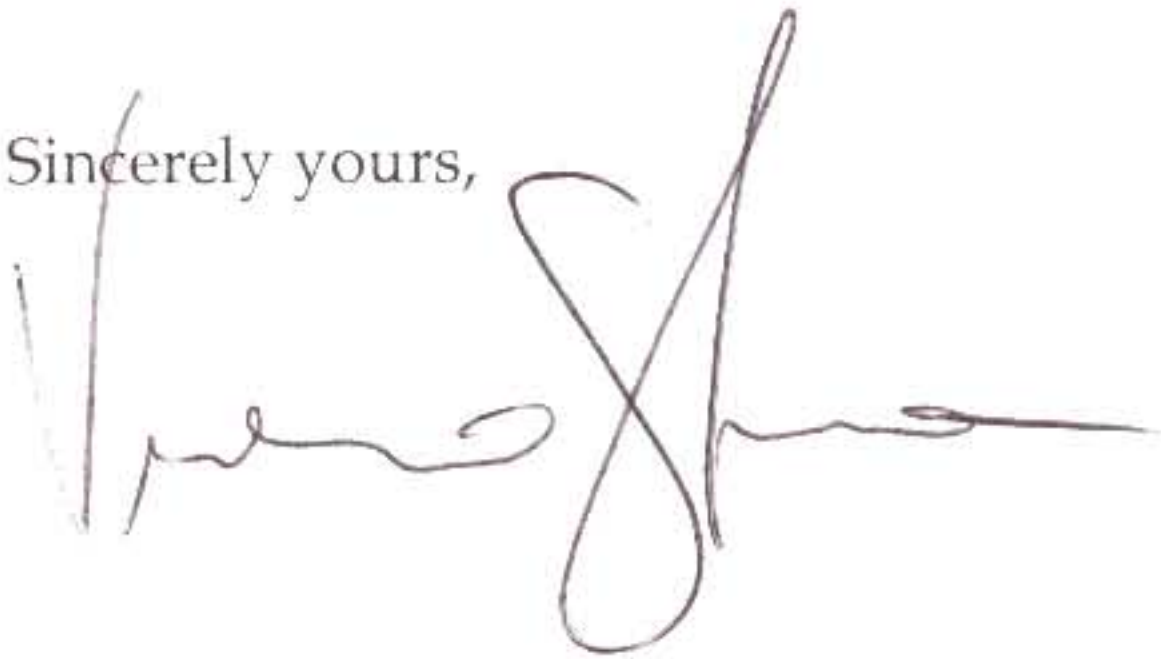
- 2) The reviewer 03552056 suggested discussing some references (HERACLES TRIAL, ICECREAM TRIAL, Dienstmann R et al. Cancer Disco 2015; 5: 598-609) and including some tables that recapitulate the clinical trials and agents in research to overcome resistance to anti-EGFR.
- 2) We have provided the information and table required.

- 3) The reviewer 02531403 suggested elucidating the inclusion and exclusion criteria of the studies discussed. He/She also suggested to define "targeted medicine" rather than "precision medicine" in the introduction and to quote studies about the role of chemoprevention and diet in the setting of EGFR mutations
- 3) We have elucidated the inclusion and exclusion criteria of the studies discussed and we have changed "target medicine" instead of precision medicine. However, we did not find studies about the role of chemoprevention and diet in the setting of EGFR mutations.

We hope that you will find the revised manuscript to be suitable for publication in World Journal of Gastroenterology.

Thank you again for publishing our manuscript in the World Journal of Gastroenterology.

Sincerely yours,

A handwritten signature in dark ink, appearing to read 'Vincenzo Sforza', with a large, stylized initial 'V' and a long horizontal stroke at the end.

Vincenzo Sforza, MD

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