

August 11th, 2021 University of Chile Faculty of Medicine Basic and Clinical Oncology Department

Lian-Sheng Ma Company Editor-in-Chief World Journal of Gastrointestinal Oncology

This letter is to submit the revision of the manuscript entitled "Consensus molecular subtypes of colorectal cancer in clinical practice: A translational approach" to be considered for publication in the Journal as a review article, for the invitation received with the NO: 69305.

I would like to provide a detailed response to each of the comments made by the manuscript reviewers. All this changes are highlighted within the text.

## **Reviewer #1:**

1.1 The following minor issues should be resolved: 1. Abstract: "for metastasis CRC" -> metastatic

**Response:** It has been decided to accept the suggestion.

1.2 [...] Abstract: "oncology routine practice" -> routine oncology care (?) **Response:** It has been decided to accept the suggestion.

1.3 Introduction: GLOBOCAN 2020 data just came out recently (DOI 10.3322/caac.21660), I recommend to update the first citation with that.

**Response:** We will update the reference; the text is maintained because of the sum of colon and rectal cancer continues to be the second cause of death worldwide.

1.4 [...] "It is associated with infiltration and immune activation" -> It is associated with lymphocyte infiltration and immune activation

**Response:** The sentence is rewritten with the reviewer's suggestion

## 1.5 [...] TGF-B -> TGF- $\beta$ (?)

**Response:** It has been decided to accept the suggestion.

1.6 [...] "and better OS in CM2 (29.0 months), whereas CM3 (18.6 months) and" -> CMS2 and CMS3

**Response:** The sentence is rewritten with the reviewer's suggestion

1.7 [...] fusion protein with successful results in murine CRC models. This treatment has demonstrated an anti-tumor response by TGF $\beta$  and PD-L1 immunosuppressive pathways with successful results in murine CRC models[46,47]." Do not replicate the same information.

**Response:** The sentence is rewritten with the reviewer's suggestion, erasing duplicated sentences.

We hope that our responses and adjustments to the manuscript will be liked by the reviewers and the Editorial Board

I look forward to hearing from you at your earliest convenience.

Sincerely yours,

## Jaime González-Montero MD PhD

Corresponding Author Basic and Clinical Oncology Department Faculty of Medicine, University of Chile. Independencia 1027, Santiago, Chile. Phone: +56999367446 E-mail: jagonzalez@ug.uchile.cl