

28/2/2020

To

Jin-Zhou Tang

Science Editor, Editorial Office

Baishideng Publishing Group Inc

Dear Editor,

On behalf of my coauthors I take this opportunity to thank you for the suggestions on the manuscript (Manuscript NO: 53580) titled 'Clinicopathological differences and correlations between right and left colon cancer'.

I am pleased to revise and submit the updated manuscript for consideration of publication as an 'Original Article' in World Journal of Clinical Cases.

We have incorporated the suggestions as detailed below. We hope that with these revisions, the manuscript will be deemed acceptable for publication.

Kindly accept minor changes regarding names and affiliations of authors.

I look forward to hearing from you.

Kind Regards

Ioannis Kalantzis

No	Reviewers Comment	Author's Response	Changes made in manuscript
Reviewer#1			
1	First of all the study suffers from an obvious selection bias.	Kindly accept our opinion that there is no selection bias to the present study since during the 2.5 year recruitment period we addressed an invitation to participate in our study to all colon cancer patients who received chemotherapy regimens in the Oncology Center. Of those patients, all patients who fulfilled the inclusion criteria were included in the study	Page 8,9 Study population and data collection
2	Randomly selected patients?	Regarding the nature of the study and the randomization of the included patients, we agree with your comment. The present study is an observational and not a randomized study. As such, we selected the patients on the basis of the location of the primary tumor and no further randomization was performed based on treatment.	Page 8,9 Study population and data collection

		Therapeutic regimens were predetermined according to RAS mutation status	
3	Which are the criterias? Who select them and how they exclude patients?	Inclusion criteria were the presence of histologically confirmed colon cancer, the patient's signed informed consent and the presence of updated medical records.	NA
4	It should be a motivation how they managed with time to recurrence as progerssion to stage IV?	Your comment about the time to recurrence is not clearly understood. All initial stage I-III patients who progressed to stage IV initiated immediately first line chemotherapy treatment	NA
5	Why they include rectum tumors if the disase in rectum and colon are different diseases?	Our decision to include rectal tumors along with left colon cancer tumors is supported by most of the already published studies (Clinicopathological and molecular differences in colorectal cancer according to location. Hsu YL, Lin CC, Jiang JK et al. Int J Biol Markers. 2019 Mar;34(1):47-53, Are there two sides to colorectal cancer? Iacopetta B. Int J Cancer. 2002 Oct 10;101(5):403-8, The	NA

		<p>significance of microsatellite instability in colorectal cancer after controlling for clinicopathological factors. Sanghee Kang, Younghyun Na, Sung Yup Joung et al. Medicine (Baltimore). 2018 Mar; 97(9), Primary tumor sidedness has an impact on prognosis and treatment outcome in metastatic colorectal cancer: results from two randomized first-line panitumumab studies. N. Boeckx, R. Koukakis, K. Op de Beeck et al. Ann Oncol. 2017 Aug; 28(8): 1862–1868) since they share common embryological origin, arterial supply, microbial populations and exposure to various toxic substances and bile acids, as well as the same molecular characteristics and tumorigenic pathways</p>	
6	How was the deal with missing variables in multivariable model?	For the purpose of the multivariate analysis no missing values were imputed.	Page 10 Statistical analysis
Reviewer#2			
1	Abbreviations (such	Appropriate corrections	Page

	as “OR”, “MAPK”, “MSI”..., etc) should be with its full name when it firstly appears in the paper.	regarding abbreviations will be performed as suggested.	4,5,8,14,15,18,
2	The paper presents some typing errors that have to be amended.	Thank you very much for this comment. It would be highly appreciated if the typing errors in our article could be indicated. We kindly remind that the manuscript has been edited and received an editing certificate by American Journal Experts.	NA