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SPECIFIC COMMENTS TO AUTHORS

This is an adequate review paper, but there are some minor revisions that are minimally necessary. First, the writing needs editing, for style, punctuation, capitalization, etc. Second, when discussing in the Introduction the various signaling pathways of importance for colorectal cancer, there needs to be some mention about how these pathways are becoming deregulated to initiate disease. For example, discussing the normal function of the canonical Wnt signaling pathway is insufficient in the context of colorectal cancer, since most cases of that disease are initiated by mutations (chiefly APC, but also beta-catenin and other minor mutations) that deregulate the signaling and are potential therapeutic targets. Third, with the analysis of various signaling pathways in the Introduction, there is then very little further discussion on how these can be targeted for therapy. For example, there is significant literature about targeting Wnt signaling. It may be helpful to at least briefly list examples of actual or proposed targeting of relevant signaling pathways. One does not expect a comprehensive discussion in what is a brief review, but at least a mention would be relevant and informative.

Answer:

As per the reviewer's first comment, the manuscript has been duly edited for style, punctuation and capitalization.

The deregulation and mutations which occur in all signaling pathways and could be exploited for targeting CRC have been added in the introductory part.

Relevant studies in which various pathways are targeted have also been discussed in the introduction as per the comment raised by the reviewer.