

Reviewer 1

In this interesting experimental study, the authors aimed to evaluate the diagnostic value of circular free DNA for CRC. Endoscopy combined with tissue biopsy is the gold standard for early diagnosis of CRC at present, but there are some disadvantages such as cumbersome operation, poor compliance and invasion. It is urgent to find a minimally invasive or non-invasive, sensitive and accurate early diagnosis method. Early detection of CRC is urgently needed in clinical practice. Commonly biomarker and circular free DNA may have potential diagnostic value for CRC. They concluded that circular free CAMK1D may be potential diagnostic biomarker for colorectal cancer detection. The study is well designed and the manuscript is appropriately written. I congratulate the authors for their successful work. Language polishing is needed to make best sense of reading.

Answer

Thanks very much for the reviewer work. And we have made revision in the language for better reading for readers.

Reviewer 2

1- Core tip section: This part writes too concise, and I suggest to summarize the methods, results and conclusions of this study.

2- Methods section: I suggest to improve the subjects enrollment algorithm. Also, I suggest to describe how many subjects are enrolled in the study sample.

3- Table section: There are two 'CRC' and 'HC' in Table 1. Although it is possible to know in the text that one group was enrolled in the establishing model and the other was included separately to validate the model, I suggest it to be stated in the table alone.

4- Discussion section: Although there is much discussion about early diagnosis of CRC, there is less discussion about the results of this study and more discussion about the results of the study needs to be added.

Answer

1. We have revised the core tip section to be more related with the topic of this study.

2. We have added the subject enrollment and added the sample size in the methods section.

3. In the table section, we have revised the CRC and HC training and validation for better reading.

4. Thanks, we have added the discussion about CAMK1D in our results in the Discussion section.