



**Baishideng
Publishing
Group**

7041 Koll Center Parkway, Suite
160, Pleasanton, CA 94566, USA
Telephone: +1-925-399-1568
E-mail: bpgoffice@wjgnet.com
https://www.wjgnet.com

Reviewer#1

SPECIFIC COMMENTS TO AUTHORS

Hypermethylation in the promoter region is the main mechanism of downregulation or silencing of RNF180 expression in tumors. The primary screening methods for GC in the clinic are upper gastrointestinal X-ray examination, gastrointestinal endoscopy, Helicobacter pylori antibody detection and plasma pepsinogen detection. ddPCR is a novel absolute quantitative technique that divides the reaction system into thousands of units. In this study, the authors used ddPCR to quantify the methylation level of the RN180 gene. The study is very well designed and the manuscript is very well written. The methods are described in detail, the results are well displayed.

Minor comments: 1. Some minor editing is required. 2. Table 1 should be re-arranged. 3. References should be edited and updated.

Answer: Thanks for the comments. We have carefully checked the editing and updated the references. In addition, the Table 1 was also ra-arranged.

Reviewer#2

SPECIFIC COMMENTS TO AUTHORS

In this study, methylation level of the RN180 gene was quantified by droplet digital PCR. The study is well performed, and the results are interesting. The reviewer recommends to accept this study after a minor editing.

Answer: Thanks for the positive comments on our study.

Reviewer#3

SPECIFIC COMMENTS TO AUTHORS

RNF180 affects many important physiological processes in vivo, including cell growth, differentiation, and tumorigenesis. RNF180 is expressed at low levels in gastric cancer tissue samples, and it is particularly crucial to the gastric cancer pathological stage and overall survival of patients. This is an interesting study about the methylation level of the



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RN180 gene in in gastric cancer. The study is very interesting. The authors should take attention about the abbreviations and some spelling mistakes. Thank you.

Answer: Thanks. We have checked the abbreviation and spelling mistakes.