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ANSWERING REVIEWERS

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name 22782-Revised manuscript.doc).

Title: Molecular alterations in gastric cancer with special reference to the early-onset subtype

Authors: Małgorzata Skierucha, Anya NA Milne, George Johan A Offerhaus, Wojciech P Polkowski, Ryszard Maciejewski and Robert Sitarz

Name of Journal: *World Journal of Gastroenterology*

Column: Review

Manuscript NO.: 22782

The manuscript has been improved according to the suggestions of reviewers:

- 1 Format has been updated
- 2 Revision has been made according to the suggestions of the reviewer
- 3 References has been double-checked and they are not repeated. Twelve references have been deleted and nine added. References and typesetting have been corrected.

ANSWERS TO REVIEWERS COMMENTS

1. Reviewer 03017455

Comment: *In this manuscript, the authors provided a summary of published data concerning the gastric cancer with special reference to early onset subtype. Minor comments- 1. The title is "Molecular alterations in gastric cancer with special reference to early onset subtype." I suggest that authors put more attention on the section "Early Onset Gastric Carcinoma" and cut down the other section. 2. In Figure 1, the authors mentioned 4 types of gastric cancers. However, authors didn't discuss gastric stump cancer in the main body part. 3. The references should be update.*

Response:

- 1) We have extended the section about Early Onset Gastric Carcinoma and divided it into two separate parts: 'Early Onset Gastric Carcinoma' at the beginning and 'Early Onset Gastric Carcinoma's Distinctive features' at the end of the manuscript. The rest sections have been shortened, some parts have been deleted.

2) Gastric stump cancer is a clinical diagnosis used for all cases of GC in the setting of the postoperative scar. Differently, early onset, conventional and hereditary GC have their distinctive pathological features. We mark in the text that the figure presents the alternative division which encompasses the subgroups which may overlap.

3) The references have been updated.

2. Reviewer 03270452

Comment: The author described the gastric cancer with special reference to early onset subtype. It was a rare subtype, and described the molecular mechanism including SNP, CIN, MSI, Epigenetic alterations, Somatic gene mutations. But There were some parts should be improved. the section "Early Onset Gastric Carcinoma" we suggested should be at the beginning not the last. and the paper should cite some recently references. and some language should be improved.

Response: We have extended the section about Early Onset Gastric Carcinoma and divided it into two separate parts: 'Early Onset Gastric Carcinoma' at the beginning and 'Early Onset Gastric Carcinoma's Distinctive features' at the end of the manuscript. The references have been updated and enriched with nine newer ones. The manuscript has been doubled checked with special attention to the language correctness and it has been certified by biomedical editing company.

3. Reviewer 02841708

Comment: In the review, authors provides a summary of so far published data concerning the molecular characteristics of GC. There is significance of this paper to provide information about carcinogenesis and development of gastric cancer. The content was comprehensive, and structure of the paper is well organized. However, I consider the paper is a little too long, and I suggest authors to revise it

Response: The paper has been revised and reorganized. The part about Early Onset Gastric Carcinoma has been divided into two sections at the beginning and at the end of the manuscript. The other parts have been shortened, some parts and references have been deleted.

4. Reviewer 00068458

Comment: The authors have reviewed good studies for "Molecular alterations in gastric cancer" and have submitted a well-written manuscript. - minor comment; 1. In early onset gastric carcinoma section, the authors reviewed EOGC distinctive features (Milne et al., 2007). Please give more detailed explanation about the difference in molecular characteristics between early onset gastric carcinoma and sporadic gastric cancer.

Response: We have extended the section about Early Onset Gastric Carcinoma and divided it into two separate parts: 'Early Onset Gastric Carcinoma' at the beginning and



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'Early Onset Gastric Carcinoma's Distinctive features' at the end of the manuscript.
The references have been updated and enriched with nine newer ones.

Thank you again for publishing our manuscript in the World Journal of Gastroenterology.

Sincerely yours,

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