

ANSWERING REVIEWERS



Nov 11, 2013

Dear Editor,

Please find enclosed the edited manuscript in Word format (file name: 5964-Review.doc).

Title: Accumulation of aberrant DNA methylation during colorectal cancer development

Author: Eiji Sakai, Atushi Nakajima, Atsushi Kaneda

Name of Journal: *World Journal of Gastroenterology*

ESPS Manuscript NO: 5964

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 reviewers.

(1) Reviewer 00503405:

"In the review article of Sakai et al. the authors aimed to review the role of aberrant DNA methylation in colorectal carcinogenesis. The review is absolutely correct, the authors logically summarized and combined the genetic and epigenetic alterations in CRC pathogenesis, moreover systematically discussed the different molecular pathways of CRC. The style is also easy-to-read. I suggest to accept the manuscript as it is for publication in WJG. Congratulations for the authors!"

--WE thank the reviewer for careful reading and appreciating our manuscript.

(2) Reviewer 00477066:

"This is an extensive and well-written review on the role of DNA methylation in colorectal carcinogenesis discussing the potential usefulness of DNA methylation as surrogate biomarkers for diagnosis, prognosis and therapy."

--WE thank the reviewer for careful reading and appreciating our manuscript.

(3) Reviewer 00503404:

"This is a high quality, comprehensive review paper. Comments; 1. Authors could add 2 or 3 Tables to summarize the major findings of the most important studies representing the different CRC development pathways (serrated, adenoma-carcinoma-do novo). This would certainly increase the readability of the paper."

--WE thank the reviewer for careful reading of our manuscript and providing helpful comments to improve our manuscript. We added two tables. Table 1 shows the most important studies on colorectal carcinogenic pathways and genetic alterations, including adenoma-carcinoma sequence, serrated pathway, and de novo pathway. Table 2 shows the most important reports on colorectal cancer classification by methylation information. We agree to the reviewer that these tables will increase the readability of our paper.

3 References and typesetting were corrected.

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

Sincerely yours,
Atsushi

A handwritten signature in black ink, appearing to be 'Atsushi Kaneda', written in a cursive style.

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