

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45711

Title: Functional role of long non-coding RNA CAS 9/miR-140-5p/CEMIP axis in colorectal cancer progression in vitro

Reviewer's code: 00070916

Reviewer's country: Germany

Science editor: Ruo-Yu Ma

Date sent for review: 2019-02-12

Date reviewed: 2019-02-12

Review time: 7 Hours

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input checked="" type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language	(High priority)	<input type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	polishing	<input checked="" type="checkbox"/> Accept	<input checked="" type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
<input type="checkbox"/> Grade E: Do not	language polishing	<input type="checkbox"/> Minor revision	topic of the manuscript:
publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input type="checkbox"/> Advanced
		<input type="checkbox"/> Rejection	<input checked="" type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

This study is in general well planned and performed. I have only one major suggestion and a few minor comments to improve the manuscript. Major: it would be helpful to check the correlation between important molecular features (subtype, BRAF, KRAS



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and/or P53 mutations, MSI Status CIMP Status) and CASC9 and CEMIP expression.
Minor: the term chemotherapy is not suited to describe a targetting of CEMIP or CASC9.
"Frozen in liquid nitrogen at -80°C" in materials and methods is incorrect.

INITIAL REVIEW OF THE MANUSCRIPT

Google Search:

- ☐ The same title
- ☐ Duplicate publication
- ☐ Plagiarism
- ☐ No

BPG Search:

- ☐ The same title
- ☐ Duplicate publication
- ☐ Plagiarism
- ☐ No

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45711

Title: Functional role of long non-coding RNA CAS 9/miR-140-5p/CEMIP axis in colorectal cancer progression in vitro

Reviewer's code: 03459541

Reviewer's country: Spain

Science editor: Ruo-Yu Ma

Date sent for review: 2019-01-29

Date reviewed: 2019-02-13

Review time: 13 Hours, 15 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input checked="" type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good	polishing	<input checked="" type="checkbox"/> Accept	<input type="checkbox"/> Onymous
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publish	<input type="checkbox"/> Grade D: Rejection	<input type="checkbox"/> Major revision	<input checked="" type="checkbox"/> Advanced
		<input type="checkbox"/> Rejection	<input type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

Wang and co-workers examine the mechanism by which the long non-coding RNA CASC19 regulates proliferation and metastasis in CCR. They found that CASC19 positively regulated CEMIP expression and epithelial-to-mesenchymal transition



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through targeting of miR-140-5p. The state-of-the art is adequate as well as the methodology used. The conclusions are in concordance with the results obtained. The paper can be published in its actual format.

INITIAL REVIEW OF THE MANUSCRIPT

Google Search:

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- ☒ No

BPG Search:

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- ☒ No

PEER-REVIEW REPORT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 45711

Title: Functional role of long non-coding RNA CAS 9/miR-140-5p/CEMIP axis in colorectal cancer progression in vitro

Reviewer's code: 02534486

Reviewer's country: Italy

Science editor: Ruo-Yu Ma

Date sent for review: 2019-02-12

Date reviewed: 2019-02-18

Review time: 12 Hours, 6 Days

SCIENTIFIC QUALITY	LANGUAGE QUALITY	CONCLUSION	PEER-REVIEWER STATEMENTS
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	<input type="checkbox"/> Accept	Peer-Review:
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of	(General priority)	Peer-reviewer's expertise on the
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			<input checked="" type="checkbox"/> No

SPECIFIC COMMENTS TO AUTHORS

Some of the methods is described with some inaccuracies. For example: - Paragraph "Wound-healing and transwell assay": How has been the wound healing assay evaluated and quantified? The Authors must describe this detail - Paragraph

“Immunohistochemistry assay”, pg 6, lanes 17-29: The method is described in a confusing manner, since the English is not fluent. I suggest to modify the text as following: “The Immunohistochemistry assay determined CEMIP expression in CRC tissues and adjacent normal colon mucosa tissues as described in our previous study [37]. Briefly, tissue samples were fixed in 4% paraformaldehyde, embedded in paraffin and sliced into 4 µm sections. Dehydration was carried out with xylene and a gradient of ethanol solution before the endogenous peroxidase blocking with a 3% H₂O₂ solution. The antigen retrieval was performed with heated sodium citrate solution (92°C–95°C, 10 nM, pH 6.0) for 5 minutes and 1% goat serum was added to the sections at room temperature for 2 minutes to block the sections. Rabbit anti-human CEMIP antibody (1:150; CST, USA) was applied onto the sections and left overnight to incubate at 4°C. After incubation with the secondary antibody 1h at RT, freshly prepared diaminobenzidine was then added to the sections, that were then stained with hematoxylin. A light microscope (Leica Microsystems, Wetzlar, Germany) was used to visualize slides and brown particles in the cytoplasm or cytomembranes were taken to indicate positive staining.”

3. RESULTS. The “Results” section is not titled. Some parts of the Results are not described accurately. Specifically:

- Paragraph “Effect of DNA incubation on colonosphere-forming capacity of HT29 cells”: Pg. 6, Lane 46: “Concurrently, the CASC19 expression in CRC cell lines was markedly elevated....” The term “Concurrently” is not appropriate: “Consistently” or “Coherently” should be more adequate. Pg 7, lanes 1-2: “For further in vitro studies, HCT-116 and SW480 cell lines were used...” I suggest to replace with: “and this was confirmed for all the four cell line used (HCT-116 and SW480 SW620 and LOVO cells)”. Pg. 7 lane 5: “based on whether the patient had lymph node or liver metastasis...” based on whether the patient had or not lymph node or liver metastasis”
- Table 1: In Table 1 TNM stages are not legible
- Figure 1E: In Figure 6E it should be an error in the legend of the bar graph,

since the nuclear marker U6 seems to be predominantly in the cytosol (about 80% in the cytosol vs about 20% in the nucleus), while that cytoplasmic marker GAPDH seems to be predominantly in the nucleus (about 15% in the cytosol vs about 85% in the nucleus)... Check this point accurately and correct it - Figure 2D: The % of early and late apoptotic cells reported in Figure 2D are not well legible: increase the font size - Paragraph "CASC19 promoted CRC cell migration and invasion in vitro": Pg. 7 lane 46 "percent wound closure (%)". How has been calculated this %? See also comments to MM - Paragraph "CASC19 promoted CEMIP expression by targeting miR-140-5p", pg. 8 lane 34: "A luciferase reporter assay was carried out to determine if miR-140-5p could directly target miR-140-5p". Perhaps it should be "A luciferase reporter assay was carried out to determine if CASC19 could directly target miR-140-5p." - Paragraph "MiR-140-5p reverses the promoting effect of CASC19 on the proliferation and metastasis of CRC cells": It is not proper to use the term "metastasis" in an in vitro systems... "metastasizing ability" should be more appropriate. This should be also modified in other points of the manuscript (same paragraph, pg. 9 lane 16; INTRODUCTION, pg.3 lane 33; CONCLUSION, pg. 9 lane 38) 4. DISCUSSION - pg. 9 lane 29: "Currently, in our study,..." should be: "In our study,..." - pg. 9 lane 31: "...research results..." should be only "...results...": "research" is redundant - pg.10 lane 8: "...there was an endogenous reaction between these two molecules..." should be "...there was an endogenous interaction between these two molecules..."

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[Y] No

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