



## PEER-REVIEW REPORT

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 54532

**Title:** TB XR1 induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

**Reviewer's code:** 02729532

**Position:** Editorial Board

**Academic degree:** MBBS, MD

**Professional title:** Associate Professor

**Reviewer's country:** India

**Author's country:** China

**Manuscript submission date:** 2020-02-03

**Reviewer chosen by:** AI Technique

**Reviewer accepted review:** 2020-02-04 09:15

**Reviewer performed review:** 2020-02-04 12:25

**Review time:** 3 Hours

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 type="checkbox"/> Grade B: Minor language	(High priority)	<input type="checkbox"/> Anonymous
<input 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
<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 type="checkbox"/> General
			<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input type="checkbox"/> No



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#### **SPECIFIC COMMENTS TO AUTHORS**

Good work. Congratulations. The authors have evaluated the role of TBL1XR1 in pancreatic cancers. By tissue assay and in vivo tests, they found that the receptor is associated with tumour progression as well as worse prognosis in pancreatic cancers.

#### **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:** 54532

**Title:** TB XR1 induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

**Reviewer's code:** 00068723

**Position:** Editorial Board

**Academic degree:** MD, PhD

**Professional title:** Doctor, Occupational Physician

**Reviewer's country:** Japan

**Author's country:** China

**Manuscript submission date:** 2020-02-03

**Reviewer chosen by:** Le Zhang

**Reviewer accepted review:** 2020-03-14 04:48

**Reviewer performed review:** 2020-03-14 05:16

**Review time:** 1 Hour

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 type="checkbox"/> Grade B: Minor language	(High priority)	<input type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	polishing	<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input checked="" type="checkbox"/> Grade D: Fair	<input checked="" 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 checked="" 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



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## **SPECIFIC COMMENTS TO AUTHORS**

The authors investigated TBL1XR1 in pancreatic cancer with clinic-pathological data, and cell experiments. The results were rationale, but construction of the manuscript was immature. The logical flow to TBL1XR1 was not fully clear. How did the authors focus TBL1XR1 remained unclear. How was informed consent was obtained? Materials and Methods was immature. This part should be revised so that the other researchers could repeat. Methods of immunostaining was absent. How was the primary antibody obtained? Intensity of TBL1XR1 positivity (0-3) should be shown as representative photos. Did “1640 medium” mean “RPMI1640”? Cell transfection. The amount of nucleic acid was not clear. Lentivirus-mediated RNA interference. Did the authors construct lentivirus vector for RNAi? If so, the transduction methods should be written. Was lipofectamine necessary for lentivirus transduction? What was CCK-8? Was it commercially available? Company names and locations of antibodies were absent. Table 1. TNM staging correlated with positivity of TBL1XR1. But tumor sized did not. Were there any speculations to this discrepancy? Figure 1B. Both adjacent and PCa tissues look accumulation of lymphocytes. Were there any other photos? For example, adjacent tissues include acinus, islets, and pancreatic duct. Pancreatic cancer tissues include cancer cells.

## **INITIAL REVIEW OF THE MANUSCRIPT**

### *Google Search:*

- The same title
- Duplicate publication
- Plagiarism
- No



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***BPG Search:***

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- No



**PEER-REVIEW REPORT**

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 54532

**Title:** TB YXR1 induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

**Reviewer's code:** 02440884

**Position:** Editorial Board

**Academic degree:** MD

**Professional title:** Professor

**Reviewer's country:** Germany

**Author's country:** China

**Manuscript submission date:** 2020-02-03

**Reviewer chosen by:** Le Zhang

**Reviewer accepted review:** 2020-03-17 08:18

**Reviewer performed review:** 2020-03-17 08:44

**Review time:** 1 Hour

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
<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 checked="" 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



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#### **SPECIFIC COMMENTS TO AUTHORS**

In the experimental study TBL1XR1 was investigated in cellular models of pancreatic cancer using several molecular and morphological techniques. A correlation between TBL1XR1 expression and worse prognosis of pancreatic cancer was found. As a molecular mechanism of the tumor promoting TBL1XR1 effect, the authors give some evidence that TBL1XR1 is able to regulate the PI3K/AKT pathway. Comments 1. The types of pancreatic carcinoma should be clearly addressed according to the WHO. 2. TABLES 1 through 3: The headline 'Histopathological subtypes' is not correct. The authors give not the entities, but the grading and it comes not clear the grading of which type of pancreatic carcinoma. 3. A two-based category for grading should be used: low and high grade. 4. Figure 1 B: the TBL1XR1 expression is shown in lymphocytes and not in tumor cells. This point is of high importance. The source of TBL1XR1 should be identified in the tissues and better correlated to the findings in cell culture. How does the authors verify that cancer cell but not lymphocytes are under investigation?

#### **INITIAL REVIEW OF THE MANUSCRIPT**

##### ***Google Search:***

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##### ***BPG Search:***

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- Duplicate publication



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[ ] Plagiarism

[ Y ] No



**PEER-REVIEW REPORT**

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 54532

**Title:** TB YXR1 induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

**Reviewer's code:** 00503623

**Position:** Editorial Board

**Academic degree:** MD, PhD

**Professional title:** Professor

**Reviewer's country:** United States

**Author's country:** China

**Manuscript submission date:** 2020-02-03

**Reviewer chosen by:** Le Zhang

**Reviewer accepted review:** 2020-03-17 14:16

**Reviewer performed review:** 2020-03-17 17:32

**Review time:** 3 Hours

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 checked="" type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input type="checkbox"/> Grade C: Good	polishing	<input checked="" 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
<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 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



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#### **SPECIFIC COMMENTS TO AUTHORS**

This manuscript, 4532, reports the results of studies on the expression of TBL1XR1 and its function and the mechanism of action in pancreatic carcinoma (PCa) development. Based on the obtained the conclusion is that the patients with TBL1XR1-positive tumors has much worse overall survival rate than those with TBL1XR1-negative tumors. Moreover, it is shown that TBL1XR1 promotes PCa tumor cell growth through PI3K/AKT signaling pathway. While the presented results certainly support the reached conclusion, it should be noted that the role of AKT in cNOS activation through phosphorylation was not explored at all. This is a serious omission, since cNOS activation through Akt induced phosphorylation plays very important role in cell apoptosis as well as in the cellular signaling (see *Inflammopharmacology*, 18 (2010)233 - 240 and *ISRN Gastroenterology* 2011, doi.org/10.5402/2011/308727). Certainly, these papers should be incorporated into the revised paper. In addition,, the paper requires some language and grammar corrections (see Discussion "PI3K/AKT signalig pathway is participated in TBL1XR1-increase.."), instead participates...

#### **INITIAL REVIEW OF THE MANUSCRIPT**

##### ***Google Search:***

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- No

##### ***BPG Search:***

- The same title



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Duplicate publication

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No



**PEER-REVIEW REPORT**

**Name of journal:** World Journal of Gastroenterology

**Manuscript NO:** 54532

**Title:** TB XR1 induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

**Reviewer's code:** 00070310

**Position:** Editorial Board

**Academic degree:** MD, PhD

**Professional title:** Director

**Reviewer's country:** Japan

**Author's country:** China

**Manuscript submission date:** 2020-02-03

**Reviewer chosen by:** Le Zhang

**Reviewer accepted review:** 2020-03-14 09:05

**Reviewer performed review:** 2020-03-18 04:42

**Review time:** 3 Days and 19 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 type="checkbox"/> Grade B: Very good	<input type="checkbox"/> Grade B: Minor language polishing	(High priority)	<input checked="" type="checkbox"/> Anonymous
<input checked="" type="checkbox"/> Grade C: Good		<input type="checkbox"/> Accept	<input type="checkbox"/> Onymous
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	(General priority)	Peer-reviewer's expertise on the topic of the manuscript:
<input type="checkbox"/> Grade E: Do not publish	<input type="checkbox"/> Grade D: Rejection	<input checked="" type="checkbox"/> Minor revision	<input type="checkbox"/> Advanced
		<input type="checkbox"/> Major revision	<input checked="" type="checkbox"/> General
		<input type="checkbox"/> Rejection	<input type="checkbox"/> No expertise
			Conflicts-of-Interest:
			<input type="checkbox"/> Yes
			<input checked="" type="checkbox"/> No



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#### **SPECIFIC COMMENTS TO AUTHORS**

This manuscript has elucidated the function and potential mechanism of TBL1XR1 in the development of pancreatic carcinoma. This paper is well written, and will be acceptable. However, it will require some revision before publication. 1, Please impact clinical relevance in this study. How were the effects of TBL1XR1 inhibitors for the patients with pancreatic carcinoma? 2, Please summarize figures.

#### **INITIAL REVIEW OF THE MANUSCRIPT**

##### ***Google Search:***

- The same title
- Duplicate publication
- Plagiarism
- No

##### ***BPG Search:***

- The same title
- Duplicate publication
- Plagiarism
- No



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RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 54532

Title: TB X induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

Reviewer's code: 00070310

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Director

Reviewer's Country/Territory: Japan

Author's Country/Territory: China

Manuscript submission date: 2020-02-03

Reviewer chosen by: Ruo-Yu Ma

Reviewer accepted review: 2020-04-14 08:34

Reviewer performed review: 2020-04-14 08:40

Review time: 1 Hour

Scientific quality  Grade A: Excellent  Grade B: Very good  Grade C: Good

Grade D: Fair  Grade E: Do not publish

Language quality  Grade A: Priority publishing  Grade B: Minor language polishing  Grade C: A great deal of language polishing  Grade D: Rejection

Conclusion  Accept (High priority)  Accept (General priority)

Minor revision  Major revision  Rejection

Peer-reviewer statements Peer-Review:  Anonymous  Onymous

Conflicts-of-Interest:  Yes  No



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#### SPECIFIC COMMENTS TO AUTHORS

This paper is well revised, and will be acceptable. However, it will be acceptable for publication.

#### RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 54532

Title: TB YX induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

Reviewer's code: 02729532

Position: Editorial Board

Academic degree: MBBS, MD

Professional title: Associate Professor

Reviewer's Country/Territory: India

Author's Country/Territory: China

Manuscript submission date: 2020-02-03

Reviewer chosen by: Ruo-Yu Ma

Reviewer accepted review: 2020-04-16 03:44

Reviewer performed review: 2020-04-16 09:46

Review time: 6 Hours

Scientific quality  Grade A: Excellent  Grade B: Very good  Grade C: Good

Grade D: Fair  Grade E: Do not publish

Language quality  Grade A: Priority publishing  Grade B: Minor language polishing  Grade C: A great deal of language polishing  Grade D: Rejection



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Conclusion [  ] Accept (High priority) [  ] Accept (General priority)

[  ] Minor revision [  ] Major revision [  ] Rejection

Peer-reviewer statements Peer-Review: [  ] Anonymous [  ] Onymous

Conflicts-of-Interest: [  ] Yes [  ] No

#### SPECIFIC COMMENTS TO AUTHORS

Good work. However, there are some minor grammar and spelling errors, which have been commented.

#### RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 54532

Title: TB YX induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

Reviewer's code: 02440884

Position: Editorial Board

Academic degree: MD

Professional title: Professor

Reviewer's Country/Territory: Germany

Author's Country/Territory: China

Manuscript submission date: 2020-02-03

Reviewer chosen by: Ruo-Yu Ma

Reviewer accepted review: 2020-04-16 15:50

Reviewer performed review: 2020-04-16 15:58

Review time: 1 Hour

Scientific quality [  ] Grade A: Excellent [  ] Grade B: Very good [  ] Grade C:



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Good

Grade D: Fair  Grade E: Do not publish

Language quality  Grade A: Priority publishing  Grade B: Minor language  
polishing  Grade C: A great deal of language polishing  Grade D: Rejection

Conclusion  Accept (High priority)  Accept (General priority)

Minor revision  Major revision  Rejection

Peer-reviewer statements Peer-Review:  Anonymous  Onymous

Conflicts-of-Interest:  Yes  No

#### SPECIFIC COMMENTS TO AUTHORS

In the revised version of the experimental study to TBL1XR1 in pancreatic cancer the essentials were addressed and I have no further questions.

#### RE-REVIEW REPORT OF REVISED MANUSCRIPT

Name of journal: World Journal of Gastroenterology

Manuscript NO: 54532

Title: TB YX induces cells proliferation and inhibit cells apoptosis by PI3K/AKT pathway in pancreatic carcinoma

Reviewer's code: 00068723

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Doctor, Occupational Physician

Reviewer's Country/Territory: Japan

Author's Country/Territory: China

Manuscript submission date: 2020-02-03

Reviewer chosen by: Ruo-Yu Ma



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Reviewer accepted review: 2020-04-15 08:56

Reviewer performed review: 2020-04-18 01:01

Review time: 2 Days and 16 Hours

Scientific quality  Grade A: Excellent  Grade B: Very good  Grade C:  
Good

Grade D: Fair  Grade E: Do not publish

Language quality  Grade A: Priority publishing  Grade B: Minor language  
polishing  Grade C: A great deal of language polishing  Grade D: Rejection

Conclusion  Accept (High priority)  Accept (General priority)

Minor revision  Major revision  Rejection

Peer-reviewer statements Peer-Review:  Anonymous  Onymous

Conflicts-of-Interest:  Yes  No

#### SPECIFIC COMMENTS TO AUTHORS

The authors successfully addressed the concerns.