

## ESPS PEER-REVIEW REPORT

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

**ESPS manuscript NO:** 24318

**Title:** Contribution of mammalian target of rapamycin in pathophysiology of cirrhotic cardiomyopathy

**Reviewer's code:** 00052926

**Reviewer's country:** Greece

**Science editor:** Ze-Mao Gong

**Date sent for review:** 2016-01-19 09:16

**Date reviewed:** 2016-01-27 04:23

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input checked="" type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input type="checkbox"/> Grade C: Good		<input type="checkbox"/> Duplicate publication	
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade E: Poor		<input checked="" type="checkbox"/> No	<input checked="" type="checkbox"/> Minor revision
	<input type="checkbox"/> Grade D: Rejected	BPG Search:	<input type="checkbox"/> Major revision
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

## COMMENTS TO AUTHORS

In this well written and interesting basic research study, the levels of phosphorylated mTOR in cirrhotic, control and rapamycin-treated rats were quantified. An increased expression of phosphorylated-mTOR in left ventricles of cirrhotic rats compared to controls was observed. In contrast, rapamycin reversed the ECG findings of cirrhotic cardiomyopathy and decreased p-mTOR level in cirrhotic rats. Moreover, altered localization of myocardial p-mTOR protein was observed in rapamycin-treated rats with cirrhosis compared to non-treated cirrhotic rats. I have only one recommendation to the authors. Please mention in more detail and more clearly the clinical significance of your findings. In addition a minor language polishing is needed like "evidence" instead of "evidences" "survival" instead of "survivals" "may open an avenue" (please change it). "Compared to" or "in comparison with" and not "compared with". "Despite" and not "despite of".

## ESPS PEER-REVIEW REPORT

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

**ESPS manuscript NO:** 24318

**Title:** Contribution of mammalian target of rapamycin in pathophysiology of cirrhotic cardiomyopathy

**Reviewer's code:** 02942549

**Reviewer's country:** Greece

**Science editor:** Ze-Mao Gong

**Date sent for review:** 2016-01-19 09:16

**Date reviewed:** 2016-01-29 03:53

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
<input type="checkbox"/> Grade A: Excellent	<input type="checkbox"/> Grade A: Priority publishing	Google Search:	<input type="checkbox"/> Accept
<input type="checkbox"/> Grade B: Very good	<input checked="" type="checkbox"/> Grade B: Minor language polishing	<input type="checkbox"/> The same title	<input type="checkbox"/> High priority for publication
<input checked="" type="checkbox"/> Grade C: Good	<input type="checkbox"/> Grade C: A great deal of language polishing	<input type="checkbox"/> Duplicate publication	<input type="checkbox"/> Rejection
<input type="checkbox"/> Grade D: Fair	<input type="checkbox"/> Grade D: Rejected	<input type="checkbox"/> Plagiarism	<input type="checkbox"/> Minor revision
<input type="checkbox"/> Grade E: Poor		<input checked="" type="checkbox"/> No	<input type="checkbox"/> Major revision
		BPG Search:	
		<input type="checkbox"/> The same title	
		<input type="checkbox"/> Duplicate publication	
		<input type="checkbox"/> Plagiarism	
		<input checked="" type="checkbox"/> No	

## COMMENTS TO AUTHORS

Comments to the authors: This is an interesting study about the role of mTOR on the pathogenesis of cirrhotic cardiomyopathy and the potential role of rapamycin on the improvement of cardiac dysfunction. I have some minor comments to make: 1. You describe the methods of statistical analysis on page 10. After that it is useless to repeat it on every figure legend. 2. On figure 2 it is better to show us if there is a statistical significant difference, regarding the QTc, between the cirrhotic/rapa rats and cirrhotic/NS rats. 3. According to first paragraph, page 11, the figure 3 is absolutely wrong. Fig 3B must take the place of Fig 3A and 3A must take the place of 3B. Additionally, on figure 3D you mention that there is no statistical significant difference among the 4 groups. This is not true. Please correct. 4. You found that cirrhotic rats have blunted cardiac contractility and increased phosphorylated mTOR on the endothelial cells but not on cardiomyocytes of left ventricles. After the administration of rapamycin the cardiac dysfunction improved. In addition, p-mTOR on the cardiomyocytes increased comparing to p-mTOR of endothelial cells. Please explain why and how this change leads to improvement of cardiac dysfunction. 5. On the field of discussion, please give



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some information about the potential contingency between mTOR and TNF- $\alpha$ . 6. Please update your references.