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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Nephrology
ESPS manuscript NO: 22572
Title: How do kinases contribute to tonicity-dependent regulation of the transcription factor NFAT5?
Reviewer's code: 00465176
Reviewer's country: United States
Science editor: Shui Qiu
Date sent for review: 2015-09-08 13:46
Date reviewed: 2015-09-09 03:11

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
[ ] Grade A: Excellent	[ ] Grade A: Priority publishing	Google Search:	[ ] Accept
[ ] Grade B: Very good	[Y] Grade B: Minor language polishing	[ ] The same title	[ ] High priority for publication
[Y] Grade C: Good	[ ] Grade C: A great deal of language polishing	[ ] Duplicate publication	[ ] Rejection
[ ] Grade D: Fair	[ ] Grade D: Rejected	[ ] Plagiarism	[ ] Minor revision
[ ] Grade E: Poor		[Y] No	[Y] Major revision
		BPG Search:	
		[ ] The same title	
		[ ] Duplicate publication	
		[ ] Plagiarism	
		[Y] No	

COMMENTS TO AUTHORS

The clinical significance of this manuscript is unclear. The authors fail to relay the message why the reader would care about the complex mechanisms involved in this pathway. The manuscript lacks focus and is too long with too many details and text. IT can be better summarized with summary figures and tables. For example rather than listing all the details in text they can present tables where they list each kinase and then in separate columns they comment on available in vitro versus in vivo data (animals vs humans) and then another column comments about future steps or if the mechanisms are unknown. Are there any therapeutic implications?

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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Nephrology
ESPS manuscript NO: 22572
Title: How do kinases contribute to tonicity-dependent regulation of the transcription factor NFAT5?
Reviewer's code: 00503339
Reviewer's country: United States
Science editor: Shui Qiu
Date sent for review: 2015-09-08 13:46
Date reviewed: 2015-09-10 04:39

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
[ ] Grade A: Excellent	[Y] Grade A: Priority publishing	Google Search:	[ ] Accept
[Y] Grade B: Very good	[ ] Grade B: Minor language polishing	[ ] The same title	[Y] High priority for publication
[ ] Grade C: Good	[ ] Grade C: A great deal of language polishing	[ ] Duplicate publication	[ ] Rejection
[ ] Grade D: Fair	[ ] Grade D: Rejected	[ ] Plagiarism	[ ] Minor revision
[ ] Grade E: Poor		[Y] No	[ ] Major revision
		BPG Search:	
		[ ] The same title	
		[ ] Duplicate publication	
		[ ] Plagiarism	
		[Y] No	

COMMENTS TO AUTHORS

This thorough and well written study of kinases that may intensify or block the effect of transcription factor NFAT5 is of broad interest and should prompt investigators to explore the value of blocking of microvascular disease, especially in those afflicted with diabetes. A major concern, however, is the established reality that major effects in blocking progression of vasculopathy and nephropathy in induced diabetes in rodents have not been translated to clinically useful clues to therapy in human subjects with diabetes. As examples, the high promise of blocking Advanced Glycosylated Endproduct (AGE) formation in induced diabetes in rodents based on treatment with Ruboxistaurin, Alagebrium, or Pyridoxamine all failed in clinical trials in patients. As shown in Zhou's Figure 1, the number of concurrent agents that may effect induction and progression of microvasculopathy is vast and most have still to be tested in trials of induced diabetes. Thus, this limitation in accepting the promise of initial studies in rodents should be added to the finish paper to be published. Overall, the field is just opeining and should be fully studied due to its great promise.

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ESPS PEER-REVIEW REPORT

Name of journal: World Journal of Nephrology
ESPS manuscript NO: 22572
Title: How do kinases contribute to tonicity-dependent regulation of the transcription factor NFAT5?
Reviewer's code: 00102963
Reviewer's country: Brazil
Science editor: Shui Qiu
Date sent for review: 2015-09-08 13:46
Date reviewed: 2015-09-15 10:48

CLASSIFICATION	LANGUAGE EVALUATION	SCIENTIFIC MISCONDUCT	CONCLUSION
[Y] Grade A: Excellent	[Y] Grade A: Priority publishing	Google Search:	[Y] Accept
[ ] Grade B: Very good	[ ] Grade B: Minor language polishing	[ ] The same title	[ ] High priority for publication
[ ] Grade C: Good	[ ] Grade C: A great deal of language polishing	[ ] Duplicate publication	[ ] Rejection
[ ] Grade D: Fair	[ ] Grade D: Rejected	[ ] Plagiarism	[ ] Minor revision
[ ] Grade E: Poor		[Y] No	[ ] Major revision
		BPG Search:	
		[ ] The same title	
		[ ] Duplicate publication	
		[ ] Plagiarism	
		[Y] No	

COMMENTS TO AUTHORS

Excellent Basic Science Review Paper. Should Be Accepted as it Is!