



PEER-REVIEW REPORT

Name of journal: *World Journal of Diabetes*

Manuscript NO: 74740

Title: Immediate-release tofacitinib reduces insulin resistance in non-diabetic active rheumatoid arthritis patients: A single-center retrospective study

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03737141

Position: Editorial Board

Academic degree: PhD

Professional title: Professor

Reviewer's Country/Territory: Egypt

Author's Country/Territory: Taiwan

Manuscript submission date: 2022-01-14

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-01-31 16:30

Reviewer performed review: 2022-02-08 20:51

Review time: 8 Days and 4 Hours

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input type="checkbox"/> Grade A: Priority publishing <input checked="" type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
Conclusion	<input type="checkbox"/> Accept (High priority) <input type="checkbox"/> Accept (General priority) <input checked="" type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No



**Baishideng
Publishing
Group**

7041 Koll Center Parkway, Suite
160, Pleasanton, CA 94566, USA
Telephone: +1-925-399-1568
E-mail: bpgoffice@wjgnet.com
https://www.wjgnet.com

Peer-reviewer statements	Peer-Review: <input checked="" type="checkbox"/> Anonymous <input type="checkbox"/> Onymous Conflicts-of-Interest: <input type="checkbox"/> Yes <input checked="" type="checkbox"/> No
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SPECIFIC COMMENTS TO AUTHORS

The manuscript fulfillment all the required criteria stated above but the 74740-Institutional Review Board Approval Form or Document is submitted I think in Chinese language.



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Title: Immediate-release tofacitinib reduces insulin resistance in non-diabetic active rheumatoid arthritis patients: A single-center retrospective study

Provenance and peer review: Invited Manuscript; Externally peer reviewed

Peer-review model: Single blind

Reviewer's code: 03302683

Position: Editorial Board

Academic degree: MD, PhD

Professional title: Chief Physician, Director, Professor

Reviewer's Country/Territory: China

Author's Country/Territory: Taiwan

Manuscript submission date: 2022-01-14

Reviewer chosen by: AI Technique

Reviewer accepted review: 2022-02-24 02:13

Reviewer performed review: 2022-03-05 02:21

Review time: 9 Days

Scientific quality	<input type="checkbox"/> Grade A: Excellent <input checked="" type="checkbox"/> Grade B: Very good <input type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
Language quality	<input checked="" type="checkbox"/> Grade A: Priority publishing <input type="checkbox"/> Grade B: Minor language polishing <input type="checkbox"/> Grade C: A great deal of language polishing <input type="checkbox"/> Grade D: Rejection
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SPECIFIC COMMENTS TO AUTHORS

The manuscript submitted by Yang Lin, et al. has investigated the effects of 24-week tofacitinib therapy on insulin sensitivity in non-diabetic active rheumatoid arthritis patients naïve or exposed to biologic therapy. In this retrospective study, they found reduced insulin resistance was achieved in non-diabetic active RA patients following 24wk of tofacitinib therapy, suggesting JAK/STAT signaling may have the potential in treating diabetic. It's an interesting study, and I have several comments as follows.

Major issues: 1. Glucocorticoids have a great impact on insulin resistance and glucose metabolism. And even with the same daily dose, the effects of long-term use and short-term use on insulin sensitivity vary greatly. So only analyzing the daily dose of prednisolone is not enough to exclude the impact of glucocorticoids. It is suggested to analyze the total exposure of prednisone in the course of treatment. 2. In discussion, "A reduction in IR has been identified in RA patients with a normal weight but not in those with obese status under anti-TNF- α therapy[35]. Despite no identified obesity in the present investigation (all patients had BMI <27 kg/m²), there were higher BMI levels for patients without IR reduction (n = 7) when compared to those with reduced IR (n = 30) in the high-IR group of patients naïve or exposed to biologic therapy (without vs with IR reduction: 24.53 \pm 2.07 vs 22.49 \pm 1.91 kg/m², P= 0.019), reflecting an influence of increased BMI on IR." But in present study, improvement of insulin sensitivity is more obvious in high-IR group than in low-IR group. While it is known to all that higher BMI is closely associated with more severe insulin resistance. How to explain this contradiction?

Minor issues: 1. There is a mistake in legend of Figure 3, "A: Homeostatic model assessment (HOMA)-insulin resistance (IR) levels in all 30 patients at weeks 0 and 24



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E-mail: bpgoffice@wjgnet.com
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after tofacitinib (TOF) therapy ($P = 0.016$);". According to the results, 30 patients should be 26 patients with active rheumatoid arthritis exposed to biologic agents. This should be corrected.