

## PEER-REVIEW REPORT

**Name of journal:** *World Journal of Gastrointestinal Oncology*

**Manuscript NO:** 76666

**Title:** Proteomic signatures of infiltrative gastric cancer by proteomic and bioinformatic analysis

**Provenance and peer review:** Unsolicited manuscript; Externally peer reviewed

**Peer-review model:** Single blind

**Reviewer's code:** 01587889

**Position:** Editorial Board

**Academic degree:** MD, MSc, PhD

**Professional title:** Associate Professor, Consultant Physician-Scientist

**Reviewer's Country/Territory:** United States

**Author's Country/Territory:** China

**Manuscript submission date:** 2022-03-25

**Reviewer chosen by:** Dong-Mei Wang

**Reviewer accepted review:** 2022-05-11 03:58

**Reviewer performed review:** 2022-05-11 23:24

**Review time:** 19 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 Grade E: Do not publish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 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
Conclusion	<input type="checkbox"/> Accept (High priority) <input checked="" type="checkbox"/> Accept (General priority) <input type="checkbox"/> Minor revision <input type="checkbox"/> Major revision <input type="checkbox"/> Rejection
Re-review	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No

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

Stomach cancer (GC) is the fourth most prevalent malignancy worldwide and ranks third in terms of mortality and its molecular mechanism pathways is not completely understood. Proteomic methodology has the potential to provide new avenues for diagnostics and for the development of novel personalized therapeutic targets. Zhang Lihua et al. analyzed the proteomes of infiltrative gastric cancer (IGC) and normal gastric frozen tissue samples using high performance liquid chromatography tandem mass spectrometry (HPLC-MS/MS) aiming to identify the differentially expressed proteins (DEPs). Twelve pairs of IGC and adjacent normal tissues were collected, and their proteomes analyzed by high performance liquid chromatography tandem mass spectrometry (HPLC-MS/MS). The key proteins were screened and functionally annotated by gene ontology (GO) and KEGG pathway analyses. The Statistical methods and software used were appropriate. They found that the proteins involved in cell cycle regulation, DNA replication and mismatch repair, and metabolism were significantly altered in IGC, and the proteomic profile may enable the discovery of novel biomarkers. Presented representative figures included volcano map of significantly differentially expressed proteins/ signature, gene ontology and pathways are complicated but followable, particularly the Western blotting was extremely helpful. Furthermore, the proteomic signatures of IGC provide insights into the possible mechanisms underlying IGC progression, which involve DNA replication, cell cycle, mismatch repair, and energy metabolism pathways. I find this clinical study intriguing and of educational value and it may help unravel the molecular mechanisms and novel biomarkers of IGC as it lays the foundation for the discovery of potential diagnostic and prognostic markers

as well as therapeutic targets in IGC. The significant DEPs identified in this study will have to be validated in a large cohort from multiple centers to provide justification for further clinical investigation. NB: In the future, authors are advised to upload the Institutional Review Board (IRB) approval letter translated in English.

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**Peer-review model:** Single blind

**Reviewer's code:** 05755618

**Position:** Peer Reviewer

**Academic degree:** FACP, MD

**Professional title:** Doctor

**Reviewer's Country/Territory:** Japan

**Author's Country/Territory:** China

**Manuscript submission date:** 2022-03-25

**Reviewer chosen by:** Dong-Mei Wang

**Reviewer accepted review:** 2022-05-18 02:35

**Reviewer performed review:** 2022-05-23 00:10

**Review time:** 4 Days and 21 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
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Re-review	<input checked="" type="checkbox"/> Yes <input type="checkbox"/> No

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

This article described proteomic signatures of infiltrative gastric cancer (IGC). They found the top 10 up-regulated proteins (MRTO4, BOP1, PES1, WDR12, BRIX1, NOP2, POLR1C, NOC2L, MYBBP1A, and TSR1) and the top 10 down-regulated proteins (NDUFS8, NDUFS6, NDUFA8, NDUFA5, NDUFC2, NDUFB8, NDUFB5, NDUFB9, UQCRC2, and UQCRC1). The study design is well, and exciting article. However, there are some concerns about this article. 1. The authors could describe the flow chart of this study methods. In addition, the authors could present the figure of high-performance liquid chromatography-tandem mass spectrometry (HPLC-MS/MS) to understand general readers. 2. The number of pairs of IGC and adjacent normal tissues was too small to generalize the result of this study. 3, The authors could describe how this result was applied clinically.

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**Peer-review model:** Single blind

**Reviewer's code:** 05689407

**Position:** Peer Reviewer

**Academic degree:** PhD

**Professional title:** Doctor

**Reviewer's Country/Territory:** Spain

**Author's Country/Territory:** China

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**Reviewer chosen by:** Dong-Mei Wang

**Reviewer accepted review:** 2022-05-18 06:23

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Scientific quality	<input type="checkbox"/> Grade A: Excellent <input type="checkbox"/> Grade B: Very good <input checked="" type="checkbox"/> Grade C: Good <input type="checkbox"/> Grade D: Fair <input type="checkbox"/> Grade E: Do not publish
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Re-review	<input type="checkbox"/> Yes <input checked="" type="checkbox"/> No



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<b>Peer-reviewer statements</b>	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**

Institutional review board approval document is not in english. Biostatistics review certificate is not correctly signed. Institutional animal care and use commite approval document is not signed, it is only two sentence in a Word document that anyone could have written. It is important to keep the same font type and size with concordance. In some sections they change.