

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 66203

Title: Current status of non-surgical treatment of locally advanced pancreatic cancer

Reviewer's code: 05120663

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: United States

Author's Country/Territory: Greece

Manuscript submission date: 2021-03-22

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-03-23 17:05

Reviewer performed review: 2021-03-23 18:32

Review time: 1 Hour

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 Grade E: Do not publish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
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

This is an overall good review on non-surgical therapies for pancreas cancers. The topic is interesting. Following points should be improved. There are many environmental, dietary and lifestyle factors that influence the microbiome (in intestinal lumen, tissue and other body sites), immune system, carcinogenic mechanisms, and response to therapy. The authors should discuss these points; influence of those factors, eg, smoking, alcohol, diet, obesity, microbiome, etc. on tumor and the immune system. These factors may influence response to therapy in each patient differentially. There are also influences of GxE (gene-by-environment interactions). More discussion on interaction of those genetic changes and the environmental factors (mentioned as above) in relation to response to treatment is important. In these contexts research on dietary / lifestyle factors, microbiome, immunity, and molecular tissue biomarkers is needed for the future direction. The authors should discuss molecular pathological epidemiology which can investigate those factors in relation to molecular pathologies, clinical outcomes, and response to treatment. Such research can be a promising direction and improve prediction of response to therapy. Strengths and challenges of molecular pathological epidemiology discussed in Gut 2011, Annu Rev Pathol 2019, etc. should be discussed in this review.

PEER-REVIEW REPORT

Name of journal: *World Journal of Gastrointestinal Oncology*

Manuscript NO: 66203

Title: Current status of non-surgical treatment of locally advanced pancreatic cancer

Reviewer's code: 05261005

Position: Peer Reviewer

Academic degree: MD

Professional title: Doctor

Reviewer's Country/Territory: Italy

Author's Country/Territory: Greece

Manuscript submission date: 2021-03-22

Reviewer chosen by: AI Technique

Reviewer accepted review: 2021-03-23 14:41

Reviewer performed review: 2021-04-02 09:17

Review time: 9 Days and 18 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 type="checkbox"/> Yes <input checked="" type="checkbox"/> No
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

Dear Authors, thank you for sharing your article. Non-surgical treatment of locally advanced pancreatic cancer is a very interesting and complex topic, considering that approximately 30%-35% of pancreatic cancer patients present with locally advanced disease at diagnosis, but only a small proportion of them manage to go to surgery after neoadjuvant treatment. Manuscript is well written, I suggest the following modifications:

- You should consider to discuss if there are studies that investigated (or are investigating) SBRT vs “conventional” radiotherapy - In the paragraph “ABLATIVE TECHNIQUE” I suggest discussing also the role of reversible electrochemotherapy for a matter of consistency. For example: Casadei R, Ricci C, Ingaldi C, Alberici L, Di Marco M, Guido A, Minni F, Serra C. Intraoperative electrochemotherapy in locally advanced pancreatic cancer: indications, techniques and results-a single-center experience. *Updates Surg.* 2020 Dec;72(4):1089-1096. doi: 10.1007/s13304-020-00782-x. Epub 2020 May 12. PMID: 32399592.