

Reviewer Name: Anonymous

Review Date: 2023-03-22 18:13

Specific Comments To Authors: Very significant study for tumor delineation of tumor taking in account intensity factor. There are limitations like sample size, taking in account few factors only and patients being Child A and B but as the authors pointed out they intend to carry forward study increasing spectrum of factors and more severe liver disease patients.

Scientific Quality: Grade A (Excellent)

Novelty of This Manuscript: Grade A (Excellent)

Creativity or Innovation of This Manuscript: Grade B (Good)

Scientific Significance of the Conclusion in This Manuscript: Grade A (Excellent)

Language Quality: Grade B (Minor language polishing)

Conclusion: Accept (High priority)

Answer:

We appreciate the reviewer's recognition and high evaluation for our research. There were some limitations in the current study, while our study results found that liver enzymes and other indicators were related to liver background fluorescence. These indicators can be used to predict the risk of intraoperative fluorescence interference and timely adjust the medication regimen of ICG before surgery, so as to improve the value of ICG in liver surgery. In the follow-up study, we will continue to expand the sample size and conduct sub-group analysis according to different clinical characteristics and medication plans, so as to make the study results more objective and accurate. In the process of revision, we correct the obvious expressions and grammatical errors in time to make the expression more clear and concise.

Thanks again for the reviewer's kind encouragement to our research.

Reviewer Name: Anonymous

Review Date: 2023-03-26 23:46

Specific Comments To Authors: Some important papers in this field are not listed in the reference section. Please expand the reference section with important papers, for ICG dynamics during surgery.

Scientific Quality: Grade C (Good)

Novelty of This Manuscript: Grade C (Fair)

Creativity or Innovation of This Manuscript: Grade B (Good)

Scientific Significance of the Conclusion in This Manuscript: Grade B (Good)

Language Quality: Grade C (A great deal of language polishing)

Conclusion: Minor revision

Answer:

We are grateful to the reviewers for their objective and valuable comments on our article. After we reviewed our manuscript, there were references papers ([7], [11], and [17]) related to the dynamic changes of ICG during surgery. Based on the revision suggestions, we have added references [6] and [12] to the introduction section of the article, and added relevant references [18], [19], [20], [21], and [22] to the discussion section to indicate examples of the role of ICG dynamic in other operations. At the same time, we have revised the article and made corresponding language improvements to address obvious language errors and expressions. Of course, we will continue to read and find other articles related to ICG dynamics in order to deepen the understanding of ICG dynamics, to improve the quality of our subsequent research.

Thank you again for the valuable feedback from the reviewers.

Reference paper are added as follow:

- [6]. WIEGAND BD, KETTERER SG, RAPAPORT E. The use of indocyanine green for the evaluation of hepatic function and blood flow in man. *Am J Dig Dis*. 1960;5:427-436. doi:10.1007/BF02232628
- [12]. Verbeek FP, Schaafsma BE, Tummers QR, et al. Optimization of near-infrared fluorescence cholangiography for open and laparoscopic surgery. *Surg Endosc*. 2014;28(4):1076-1082. doi:10.1007/s00464-013-3305-9
- [18]. Kraft JC, Treuting PM, Ho RJY. Indocyanine green nanoparticles undergo selective lymphatic uptake, distribution and retention and enable detailed mapping of lymph vessels, nodes and abnormalities. *J Drug Target*. 2018;26(5-6):494-504. doi:10.1080/1061186X.2018.1433681
- [19]. Zhang L, Cheng M, Lin Y, et al. Ultrasound-assisted carbon nanoparticle suspension mapping versus dual tracer-guided sentinel lymph node biopsy in patients with early breast cancer (ultraCars): phase III randomized clinical trial. *Br J Surg*. 2022;109(12):1232-1238. doi:10.1093/bjs/znac311
- [20]. Müller D, Stier R, Straatman J, et al. ICG-Lymphknoten-Mapping in der Tumorchirurgie des oberen Gastrointestinaltrakts [ICG lymph node mapping in cancer surgery of the upper gastrointestinal tract]. *Chirurgie (Heidelb)*. 2022;93(10):925-933. doi:10.1007/s00104-022-01659-y
- [21]. Wiig H, Swartz MA. Interstitial fluid and lymph formation and transport: physiological regulation and roles in inflammation and cancer. *Physiol Rev*. 2012 Jul;92(3):1005-60. doi: 10.1152/physrev.00037.2011

[22]. Nanjee MN, Cooke CJ, Olszewski WL, Miller NE. Lipid and apolipoprotein concentrations in prenodal leg lymph of fasted humans. Associations with plasma concentrations in normal subjects,