



OPEN ACCESS

EDITED AND REVIEWED BY
Antonino Belfiore,
University of Catania, Italy

*CORRESPONDENCE

Pengyuan Zheng
✉ medp7123@126.com
Ihtisham Bukhari
✉ bukhari5408@gmail.com

SPECIALTY SECTION

This article was submitted to
Cancer Endocrinology,
a section of the journal
Frontiers in Endocrinology

RECEIVED 04 March 2023

ACCEPTED 23 March 2023

PUBLISHED 13 April 2023

CITATION

Mi Y, Iqbal F, Mahmood N, Bukhari I
and Zheng P (2023) Editorial: Chronology
of gastrointestinal cancers and
gastrointestinal microbiota.
Front. Endocrinol. 14:1179413.
doi: 10.3389/fendo.2023.1179413

COPYRIGHT

© 2023 Mi, Iqbal, Mahmood, Bukhari
and Zheng. This is an open-access article
distributed under the terms of the [Creative
Commons Attribution License \(CC BY\)](#). The
use, distribution or reproduction in other
forums is permitted, provided the original
author(s) and the copyright owner(s) are
credited and that the original publication in
this journal is cited, in accordance with
accepted academic practice. No use,
distribution or reproduction is permitted
which does not comply with these terms.

Editorial: Chronology of gastrointestinal cancers and gastrointestinal microbiota

Yang Mi¹, Furhan Iqbal², Nasir Mahmood^{3,4},
Ihtisham Bukhari^{1*} and Pengyuan Zheng^{1*}

¹Henan Key Laboratory of Helicobacter pylori, Microbiota and Gastrointestinal Cancers, Marshall
Medical Research Center, Fifth Affiliated Hospital of Zhengzhou University, Zhengzhou, China,

²Institute of Zoology, Bahauddin Zakariya University, Multan, Pakistan, ³Department of Biochemistry,
Human Genetics and Molecular Biology, University of Health Sciences, Lahore, Pakistan, ⁴Department
of Cell and Systems Biology, University of Toronto, Toronto, ON, Canada

KEYWORDS

chronology, gastrointestinal cancers, gut microbiota, carcinogens, cancer development
and metastasis

Editorial on the Research Topic

Chronology of gastrointestinal cancer and gastrointestinal microbiota

The term “chronology of cancer” describes a number of factors, including the changing nature of cancers over time, extending from carcinogenesis to development, progression, and metastasis (1, 2). A number of factors can lead to the development of gastrointestinal cancer, including, but not limited to, food, infections, DNA damage, environmental carcinogens, radiation, toxic chemicals, and genetic and epigenetic alterations (3–5). Various chronologies of gastrointestinal cancers have been reported that have brought awareness to the masses and helped to develop cancer prevention strategies. Analyzing tumor growth helps to estimate the time at which cancer or metastasis occurred and to predict the survival of cancer patients (2, 6–8). These chronologies vary from case to case and may even differ for the cancers derived from the same organ (2, 9). Gut microbiota maintains a healthy body and preserves symbiosis with the host immune system that generates protective responses against pathogens and regulatory pathways that sustain tolerance to commensal microbes (10, 11). Recently, the gut microbiota came into the limelight due to their association with multiple cancers, especially gastrointestinal cancers (6, 8, 10). It is already an established fact that chronic *Helicobacter pylori* infection is an essential risk factor for atrophic gastritis, intestinal metaplasia and gastric cancer (12, 13). Recently, it has been highlighted that the gut resident microbiota influences the response to cancer-related therapies (14–16). However, gaps still exist regarding the functional activity and the microbiome changes in the gut during cancer development and progression. Thus, the alterations in the gut microbiota are being monitored to assess the antitumor response of the drugs and the modulation of the intestinal immune system (17, 18). It has been indicated that dysbiosis of gut microbiota and associated metabolites contribute to carcinogenesis through multiple pathways, such as inducing inflammation, immune

dysregulation, and genetic instability (19–21). The current Research Topic aims to report the data related to the influence of gut microbiota on the development of gastrointestinal (GI) cancers (including esophageal, gastric, colorectal, liver, and pancreatic cancers) and new strategies for the prevention and treatment of GI cancers. After rigorous screening and reviewing, only four articles exploring new dimensions in this research field were included in the current topic.

Liu et al. studied the safety features of natural orifice specimen extraction (NOSE) in treating 125 patients with the sigmoid colon and upper rectal cancer. Specifically, they focused on the clinical and pathological characteristics, inflammatory and immune indicators, and postoperative complications following NOSE application. Liu et al. divided the patients into two groups based on the treatment they received, Conventional laparoscopic-assisted radical resection for CRC (CLA) and laparoscopic-assisted radical resection for CRC with NOSE (La-NOSE) and found no significant difference in post-surgery complications in both groups. Liu et al. suggested that even with a high rate of intra-abdominal bacterial load, NOSE is still safe and an alternative therapeutic option for colorectal cancer.

Cai et al. retrospectively evaluated the Regional Lymph node Metastasis (LNM) pattern to strategize the optimal surgery standpoint of lymph node distribution in 6622 patients that were diagnosed with localized well-differentiated gastroenteropancreatic neuroendocrine tumors (GEP-NETs) that were ≤ 20 mm in size. 36% of patients showed LNM after regional lymphadenectomy. Nodal involvement was observed in different GI cancers at different proportions. It was concluded that patients at a young age with big tumors and muscular invasion had higher LNM and lymphadenectomy can help to increase the overall survival of NETs patients.

Lv et al. found that acupuncture can improve gut function and enhance the microbiota's quality after chemotherapy in mice. Furthermore, a significant correlation was observed between gut microbiota, inflammation, and serum metabolites. This study opened new doors for basic and clinical research on cancer-related fatigue after cancer therapy.

Wang et al. determined the plasma level of lncRNA lnc-MyD88 in 98 hepatocellular carcinomas (HCC) patients. In addition, they analyzed the relationship of lnc-MyD88 with clinicopathological features and immune cell infiltration in the liver cancer tumor microenvironment. Notably, they found high expression of lnc-MyD88 in HCC and a positive correlation with immune function and suggested it to be used as a diagnostic marker for liver cancer.

References

1. Lote H, Spiteri I, Ermini L, Vatsiou A, Roy A, McDonald A, et al. Carbon dating cancer: defining the chronology of metastatic progression in colorectal cancer. *Ann Oncol* (2017) 28:1243–9. doi: 10.1093/annonc/mdx074
2. Murakami K, Matsubara H. Chronology of gastrointestinal cancer. *Surg Today* (2018) 48:365–70. doi: 10.1007/s00595-017-1574-y
3. Fu T, Coulter S, Yoshihara E, Oh TG, Fang S, Cayabyab F, et al. FXR regulates intestinal cancer stem cell proliferation. *Cell* (2019) 176:1098–1112.e18. doi: 10.1016/j.cell.2019.01.036
4. Nadeem MS, Kumar V, Al-Abbasi FA, Kamal MA, Anwar F. Risk of colorectal cancer in inflammatory bowel diseases. *Semin Cancer Biol* (2020) 64:51–60. doi: 10.1016/j.semcancer.2019.05.001
5. Zhao H, He M, Zhang M, Sun Q, Zeng S, Chen L, et al. Colorectal cancer, gut microbiota and traditional Chinese medicine: A systematic review. *Am J Chin Med* (2021) 49:805–28. doi: 10.1142/S0192415X21500385
6. Bordry N, Astaras C, Ongaro M, Goossens N, Frossard JL, Koessler T. Recent advances in gastrointestinal cancers. *World J Gastroenterol* (2021) 27:4493–503. doi: 10.3748/wjg.v27.i28.4493

Conclusion and prospects

Gastrointestinal (GI) cancers are ranked among the most frequently reported cancer type worldwide and rated as the third leading cause of cancer-associated deaths. Chemotherapy, immunotherapy, radiation, and surgery are treatment options for various cancers, including GI. Since the chronology of gastrointestinal cancer is not fully explored, we provide some evidence to support the advancements in clinical and basic research, especially in gastroenterology. In the current topic, we selected studies describing the different aspects of GI cancers, including gut microbiota, the role of lncRNAs, Lymph node Metastasis (LNM), and NOSE therapy. Further and advanced studies are required to explain gastrointestinal cancers' chronology and underlying mechanisms for effective treatment.

Author contributions

All authors listed have made a substantial, direct, and intellectual contribution to the work and approved it for publication.

Acknowledgments

We gratefully acknowledge the role of our host institutes in this study.

Conflict of interest

The authors declare that the research was conducted in the absence of any commercial or financial relationships that could be construed as a potential conflict of interest.

Publisher's note

All claims expressed in this article are solely those of the authors and do not necessarily represent those of their affiliated organizations, or those of the publisher, the editors and the reviewers. Any product that may be evaluated in this article, or claim that may be made by its manufacturer, is not guaranteed or endorsed by the publisher.

7. Lacourse KD, Johnston CD, Bullman S. The relationship between gastrointestinal cancers and the microbiota. *Lancet Gastroenterol Hepatol* (2021) 6:498–509. doi: 10.1016/S2468-1253(20)30362-9
8. Wang F, Song M, Lu X, Zhu X, Deng J. Gut microbes in gastrointestinal cancers. *Semin Cancer Biol* (2022) 86:967–75. doi: 10.1016/j.semcancer.2021.03.037
9. Krueger H, Mclean D, Williams D. Gastrointestinal cancers. *Prog Exp Tumor Res* (2008) 40:85–91. doi: 10.1159/000151876
10. Engstrand L, Graham DY. Microbiome and gastric cancer. *Dig Dis Sci* (2020) 65:865–73. doi: 10.1007/s10620-020-06101-z
11. Stewart OA, Wu F, Chen Y. The role of gastric microbiota in gastric cancer. *Gut Microbes* (2020) 11:1220–30. doi: 10.1080/19490976.2020.1762520
12. Wang F, Meng W, Wang B, Qiao L. Helicobacter pylori-induced gastric inflammation and gastric cancer. *Cancer Lett* (2014) 345:196–202. doi: 10.1016/j.canlet.2013.08.016
13. Guo Y, Zhang Y, Gerhard M, Gao JJ, Mejias-Luque R, Zhang L, et al. Effect of helicobacter pylori on gastrointestinal microbiota: a population-based study in linqu, a high-risk area of gastric cancer. *Gut* (2020) 69:1598–607. doi: 10.1136/gutjnl-2019-319696
14. Seeneevassen L, Bessede E, Megraud F, Lehours P, Dubus P, Varon C. Gastric cancer: Advances in carcinogenesis research and new therapeutic strategies. *Int J Mol Sci* (2021) 22(7):3418. doi: 10.3390/ijms22073418
15. Bessede E, Megraud F. Microbiota and gastric cancer. *Semin Cancer Biol* (2022) 86:11–7. doi: 10.1016/j.semcancer.2022.05.001
16. Liu B, Li Y, Suo L, Zhang W, Cao H, Wang R, et al. Characterizing microbiota and metabolomics analysis to identify candidate biomarkers in lung cancer. *Front Oncol* (2022) 12:1058436. doi: 10.3389/fonc.2022.1058436
17. Xu X, Ying J. Gut microbiota and immunotherapy. *Front Microbiol* (2022) 13:945887. doi: 10.3389/fmicb.2022.945887
18. Singhal S, Bhadana R, Jain BP, Gautam A, Pandey S, Rani V. Role of gut microbiota in tumorigenesis and antitumoral therapies: an updated review. *Biotechnol Genet Eng Rev* (2023), 1–27. doi: 10.1080/02648725.2023.2166268
19. Meng C, Bai C, Brown TD, Hood LE, Tian Q. Human gut microbiota and gastrointestinal cancer. *Genomics Proteomics Bioinf* (2018) 16:33–49. doi: 10.1016/j.gpb.2017.06.002
20. Bakhti SZ, Latifi-Navid S. Interplay and cooperation of helicobacter pylori and gut microbiota in gastric carcinogenesis. *BMC Microbiol* (2021) 21:258. doi: 10.1186/s12866-021-02315-x
21. Dai D, Yang Y, Yu J, Dang T, Qin W, Teng L, et al. Interactions between gastric microbiota and metabolites in gastric cancer. *Cell Death Dis* (2021) 12:1104. doi: 10.1038/s41419-021-04396-y