## Check for updates

## **OPEN ACCESS**

EDITED AND REVIEWED BY Catherine Sautes-Fridman, U1138 Centre de Recherche des Cordeliers (CRC), France

\*CORRESPONDENCE Jianying Zhang jzhang@utep.edu Xiangqian Guo xqguo@henu.edu.cn

SPECIALTY SECTION This article was submitted to Cancer Immunity and Immunotherapy, a section of the journal Frontiers in Oncology

RECEIVED 16 June 2022 ACCEPTED 07 July 2022 PUBLISHED 20 July 2022

#### CITATION

Zhang J, Guo X, Jin B and Zhu Q (2022) Editorial: Tumor-associated antigens and their autoantibodies, from discovering to clinical utilization. *Front. Oncol.* 12:970623. doi: 10.3389/fonc.2022.970623

## COPYRIGHT

© 2022 Zhang, Guo, Jin and Zhu. 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: Tumor-associated antigens and their autoantibodies, from discovering to clinical utilization

## Jianying Zhang<sup>1\*</sup>, Xiangqian Guo<sup>2\*</sup>, Bilian Jin<sup>3</sup> and Qing Zhu<sup>4</sup>

<sup>1</sup>Department of Biological Sciences, The University of Texas at El Paso, El Paso, TX, United States, <sup>2</sup>Department of Preventive Medicine, Institute of Biomedical Informatics, Bioinformatics Center, Henan Provincial Engineering Center for Tumor Molecular Medicine, School of Basic Medical Sciences, Academy for Advanced Interdisciplinary Studies, Henan University, Kaifeng, China, <sup>3</sup>Cancer Center, Dalian Medical University, Dalian, China, <sup>4</sup>West China Hospital, Sichuan University, Chengdu, China

#### KEYWORDS

cancer, tumor-associated antigen, autoantibody, biomarker, immunodiagnosis

## Editorial on the Research Topic

Tumor-Associated Antigens and Their Autoantibodies: From Discovering to Clinical Utilization

The detection of autoantibodies in human sera can be used as an important indicator for the diagnosis of autoimmune diseases and has been widely used in clinical research and clinical practice. Researchers in cancer field have found that various types of autoantibodies against cell self-antigens also exist in the sera from cancer patients (1, 2). These antigens related to tumorigenesis are called tumor-associated antigen (TAA). Through the study of the molecular structure and function of some human autoantigens, it is found that there are a wide variety of cellular autoantigens that induce autoantibody responses. Therefore, these cancer-associated autoantibodies might be used as biomarkers for immunodiagnosis of certain type of cancer, or as a tool to monitor therapy as well as as an indicator to predict disease prognosis (3-9). Different approach and technology, including serological analysis of recombination cDNA expression libraries (SEREX) and proteomics, have been extensively used in the identification of TAAs in cancer (10, 11). A bunch of proteins in cancer and pre-cancer conditions were identified and characterized (12-17). More recently, we have noticed that some of other approaches were also used to identify TAAs and detect anti-TAAs autoantibodies, such as whole genome derived peptide arrays and proteome microarray technology (18–20).

In recent two decades, major developments have been made in the field of research on TAAs and anti-TAAs autoantibodies, and many studies have demonstrated that serum anti-TAAs autoantibodies can be used as effective biomarkers for cancer immunodiagnosis (1–9). The diagnostic value, clinical utility, and pathogenic significance of TAAs or anti-TAAs autoantibodies are the focus of ongoing research. This special issue mainly focus on the recent studies associated with the idea and possibility that identification of TAAs and their anti-TAAs autoantibodies can be useful for cancer immunodiagnosis and cancer immunotherapies.

In this Research Topic, our guest editors have invited investigators to contribute original research articles as well as review articles which were mainly related to TAAs and anti-TAAs autoantibodies in cancer immunodiagnosis and cancer immunotherapies, and assembled the current Research Topicfor updating the recent advances in this field. In this special issue, we have received 29 submitted manuscripts, and 18 manuscripts with 139 authors have been accepted for publication. For example, a paper of Jiang et al. used a protein array technology and identify a panel of anti-TAAs autoantibodies in the early detection of lung cancer; a paper of Liu et al. used neoantigen reactive T cells combined with tomotherapy to treat a patient with advanced HCC, who reached a long time progress free survival; a paper of Qiu et al. has evaluated the diagnostic value of autoantibody against PDLIM1 for improving the detection of ovarian cancer; a study from Wang et al. has tested and validated anti-14-3-3 zeta autoantibody might be a biomarker for predicting hepatocarcinogenesis; a study from Lu et al. suggests that the combined application of PD-1-based immunotherapy and anticancer drugs has become a new expectation for clinical treatment of colorectal cancer. In addition to these original research papers, several review articles were also included in this Research Topic. For example, a review article from Zhang et al. has summarized the latest advances in the classification of immunotherapy and the process of classification, identification and synthesis of tumorspecific neoantigens, as well as their role in current cancer immunotherapy; a review article from Li et al. has provided an overview of the tumor-associated antigens and anti-TAAs autoantibodies as biomarkers in the immunodiagnosis of steosarcoma; a review article from Jin and Wang has proposed a concept of immunogenic cell death (ICD)-based cancer vaccines and summarized sources of ICD-based cancer vaccines and their challenges, which may broaden the understandings of ICD and cancer vaccines in cancer immunotherapy.

# References

1. Tan EM, Zhang JY. Autoantibodies to tumor-associated antigens: reporters from the immune system. *Immunol Rev* (2008) 222:328–40. doi: 10.1111/j.1600-065X.2008.00611.x

2. Zhang JY, Tan EM. Autoantibodies to tumor-associated antigens as diagnostic biomarkers in hepatocellular carcinoma and other solid tumors. *Expert Rev Mol Diagn* (2010) 10(3):321–8. doi: 10.1586/erm.10.12

3. Lubin R, Zalcman G, Bouchet L, Tredaniel J, Legros Y, Cazals D, et al. Serum p53 antibodies as early markers of lung cancer. *Nat Med* (1995) 1(7):701–2. doi: 10.1038/nm0795-701

4. Soussi T. p53 antibodies in the sera of patients with various types of cancer: A review. *Cancer Res* (2000) 60(7):1777–88.

5. Zhang JY, Casiano CA, Peng XX, Koziol JA, Chan EKL, Tan EM. Enhancement ot antibody detection in cancer using panel of recombinant

In summary, this Research Topic covers many important aspects in cancer immunology, especially relating to TAAs and anti-TAAs autoantibodies. This Research Topicalso includes recent advances in the basic and clinical studies relating to cancer immunodiagnosis and cancer immunotherapy. We hope that this special issue can provide useful and helpful information to investigators in this field.

# Author contributions

JZ and XG have written the manuscript and BJ and QZ have revised the manuscript. All authors contributed to the article and approved the submitted version.

## Funding

This editorial was supported by the Project of Higher Interdisciplinary Research Institute of Henan University (No. Y21008L).

# 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.

tumor-associated antigens. Cancer Epidemiol Biomarkers Prev (2003) 12(2):136-43.

8. Dai L, Ren P, Liu M, Imai H, Tan EM, Zhang JY. Using immunomic approach to enhance tumor-associated autoantibody detection in diagnosis of hepatocellular carcinoma. *Clin Immunol* (2014) 152(1-2):127–39. doi: 10.1016/j.clim.2014.03.007

9. Zhu Q, Liu M, Dai L, Ying X, Ye H, Zhou Y, et al. Using immunoproteomics to identify tumor-associated antigens (TAAs) as biomarkers in cancer immunodiagnosis. *Autoimmun Rev* (2013) 12(12):1123–8. doi: 10.1016/j.autrev.2013.06.015

<sup>6.</sup> Zhang JY, Megliorino R, Peng XX, Tan EM, Chen Y, Chan EKL. Antibody detection using tumor-associated antigen mini-array in immunodiagnosing human hepatocellular carcinoma. *J Hepatol* (2007) 46(1):107–14. doi: 10.1016/j.jhep.2006.08.010

<sup>7.</sup> Chen Y, Zhou Y, Qiu S, Wang K, Liu S, Peng X-X, et al. Autoantibodies to tumor-associated antigens combined with abnormal alpha-fetoprotein enhance immunodiagnosis of hepatocellular carcinoma. *Cancer Lett* (2010) 289(1):32–9. doi: 10.1016/j.canlet.2009.07.016

10. Chen Y-T, Gure AO, Scanlan MJ. Serological analysis of expression cDNA libraries (SEREX) - an immunoscreening technique for identifying immunogenic tumor antigens. In: GH Su, editor. *Methods in molecular medicine* (2005). Clifton, New Jersey, USA: Humana Press. p. 207–16.

11. Le Naour F, Brichory F, Beretta L, Hanash SM. Identification of tumorassociated antigens using proteomics. *Technol Cancer Res Treat* (2002) 1(4):257– 62. doi: 10.1177/153303460200100406

12. Imai H, Chan EKL, Kiyosawa K, Fu XD, Tan EM. Nuclear autoantigen with splicing factor motifs identified with antibody from hepatocellular carcinoma. *J Clin Invest* (1993) 92(5):2419–26. doi: 10.1172/JCI116848

13. Zhang JY, Chan EKL, Peng XX, Tan EM. A novel cytoplasmic protein with RNA-binding motifs is an autoantigen in human hepatocellular carcinoma. *J Exp Med* (1999) 189(7):1101–10. doi: 10.1084/jem.189.7.1101

14. Looi KS, Nakayasu ES, de Diaz RA, Tan EM, Almeida IC, Zhang JY. Using proteomic approach to identify tumor-associated antigens as markers in hepatocellular carcinoma. *J Proteome Res* (2008) 7(9):4004–12. doi: 10.1021/pr800273h

15. Zhang J, Wang K, Zhang J, Liu SS, Dai L, Zhang JY. Using proteomic approach to identify tumor-associated proteins as biomarkers in human

esophageal squamous cell carcinoma. J Proteome Res (2011) 10(6):2863-72. doi: 10.1021/pr200141c

16. Peng B, Huang X, Nakayasu ES, Petersen JR, Qiu S, Almeida IC, et al. Using immunoproteomics to identify alpha-enolase as an autoantigen in liver fibrosis. *J Proteome Res* (2013) 12(4):1789–96. doi: 10.1021/pr3011342

17. Dai L, Li J, Xing M, Sanchez TW, Casiano CA, Zhang J-Y. Using serological proteome analysis to identify serum anti-nucleophosmin 1 autoantibody as a potential biomarker in european-american and african-american patients with prostate cancer. *Prostate* (2016) 76(15):1375–86. doi: 10.1002/pros.23217

18. Yang L, Wang J, Li J, Zhang H, Guo S, Yan M, et al. Identification of serum biomarkers for gastric cancer diagnosis using a human proteome microarray. *Mol Cell Proteomics* (2016) 15(2):614–23. doi: 10.1074/mcp.M115.051250

19. Pan J, Song G, Chen D, Li Y, Liu S, Hu S, et al. Identification of serological biomarkers for early diagnosis of lung cancer using a protein array-based approach. *Mol Cell Proteomics* (2017) 16(12):2069–78. doi: 10.1074/mcp.RA117.000212

20. Yan Y, Sun N, Wang H, Kobayashi M, Ladd JJ, Long JP, et al. Whole genome-derived tiled peptide arrays detect prediagnostic autoantibody signatures in non-small-cell lung cancer. *Cancer Res* (2019) 79(7):1549–57. doi: 10.1158/0008-5472.CAN-18-1536