

Maximizing Multi-dimensional Big Data to Explore Risk and Benefit Ratio of Immunotherapy

Ying Jing¹, Yongchang Zhang², Xinying Xue, Leng Han^{4,5}

¹ Center for Intelligent Medicine Research, Greater Bay Area Institute of Precision Medicine
(Guangzhou), School of Life Sciences, Fudan University, P. R. China

²Department of Medical Oncology, Lung Cancer and Gastrointestinal Unit, Hunan cancer hospital/The
Affiliated Cancer Hospital of Xiangya School of Medicine, Central South University, P. R. China

³Department of Respiratory and Critical Care Medicine, The Affiliated Beijing Shijitan Hospital of
Capital Medical University, P. R. China

⁴Center for Epigenetics and Disease Prevention, Institute of Biosciences and Technology, Texas A&M
University, US

⁵Department of Translational Medical Sciences, College of Medicine, Texas A&M University, US

Email: jingying@ipm-gba.org.cn

Immune-checkpoint inhibitors (ICIs) have transformed patient care in oncology but are associated with a unique spectrum of organ-specific inflammatory toxicities known as immune-related adverse events (irAEs). Given the expanding use of ICIs, an increasing number of patients with cancer experience irAEs, including severe irAEs. Proper diagnosis and management of irAEs are important to optimize the quality of life and long-term outcomes of patients receiving ICIs; however, owing to the substantial heterogeneity within irAEs, and despite multicentre initiatives, performing clinical studies of these toxicities with a sufficient cohort size is challenging. Pioneering studies from the past few years have demonstrated that aggregate clinical data, real-world data (such as data on pharmacovigilance or from electronic health records) and multi-omics data are alternative tools well suited to investigating the underlying mechanisms and clinical presentations of irAEs. We have developed research strategies to utilize different sources of 'big data' for the study of irAEs. By maximizing the unique power of big data, we successfully identified biomarkers of irAE risk, characterized the effects of demographic and anthropometric factors on irAE risk and constructed landscape of toxicities of CAR-T therapy. Harnessing big data will accelerate research on irAEs and provide key insights that will improve the clinical benefits of patients receiving immunotherapies.

Keywords: immune-related adverse events; cancer immunotherapy; clinical benefits; anti-PD-1/PD-L1