

## REGULATORY INNATE LYMPHOID CELLS SUPPRESS ANTI-TUMOUR T CELLS

Sarah Q. Crome<sup>1</sup>, Linh T. Nguyen<sup>1</sup>, Sandra Lopez-Verges<sup>2</sup>, Bernard Martin<sup>1</sup>, Anca Milea<sup>1</sup>, Ramlogan Sowamber<sup>1</sup>, Jennifer Yam<sup>1</sup>, Michael Pniak<sup>1</sup>, Jessica Nie<sup>1</sup>, Pei Hua Yen<sup>1</sup>, Sarah Rachel Katz<sup>3</sup>, Marcus Bernardini<sup>3</sup>, Patricia A. Shaw<sup>1,4</sup>, Hal K. Berman<sup>1,4</sup>, Lewis L. Lanier<sup>2</sup> and Pamela S. Ohashi<sup>1</sup>

<sup>1</sup>The Campbell Family Institute for Breast Cancer Research, Princess Margaret Cancer Centre, Toronto ON. <sup>2</sup>University of California, Department of Microbiology & Immunology, San Francisco CA. <sup>3</sup>Division of Gynecologic Oncology, University Health Network, Toronto ON. <sup>4</sup>Department of Pathology, University Health Network, Toronto ON

Anti-tumour T cells are subject to multiple mechanisms of negative regulation. Recent findings that innate lymphoid cells (ILCs), including natural killer (NK) cells, regulate adaptive T cell responses led us to examine their regulatory potential in the context of cancer. We have identified a novel ILC subset that regulates the activity of tumour-infiltrating lymphocytes (TIL). Our approach allowed us to isolate human regulatory ILCs from high-grade serous ovarian tumours, define their suppressive capacity *in vitro*, and perform a comprehensive analysis of their phenotype and function. Notably, the presence of regulatory ILCs in TIL cultures correlated with impaired T cell expansion and a striking reduction in the time to disease recurrence in patients. Functional studies revealed that regulatory ILCs suppressed both CD4<sup>+</sup> and CD8<sup>+</sup> TIL expansion and cytokine production. ILCs with regulatory potential could be distinguished phenotypically from conventional NK cells and other ILCs, suggesting they may constitute a novel innate lymphocyte population. These studies demonstrate a previously unidentified cell population regulates tumour-associated T cells.