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Neutrophils Suppress Intraluminal NK Cell–Mediated Tumor Cell Clearance and Enhance Extravasation of Disseminated Carcinoma Cells ................. 630

Précis: Neutrophils facilitate the intermediate stages of the invasion–metastasis cascade by suppressing NK-cell activity and promoting extravasation of carcinoma cells.

In Vivo Genetic Screens of Patient-Derived Tumors Revealed Unexpected Frailty of the Transformed Phenotype ............ 650

Précis: PDX shRNA screens of patient-derived melanomas identified significant interpatient heterogeneity in essential, nonmutated genes.

Identification of Predictive Biomarkers for Cytokine Release Syndrome after Chimeric Antigen Receptor T-cell Therapy for Acute Lymphoblastic Leukemia ............... 664

Précis: Early cytokine measurements can accurately predict which patients will develop severe cytokine release syndrome following chimeric antigen receptor T-cell therapy.

See commentary, p. 579

ON THE COVER

Spiegel and colleagues investigated the function of neutrophils in the intermediate stages of the invasion–metastasis cascade. Splenectomy to reduce the reservoir of myeloid cells suppressed pulmonary metastases in a murine mammary carcinoma model, and was associated with neutrophilia. Further, G-CSF overexpression phenocopied the neutrophilia and metastatic potential of the cancer cells, and G-CSF–induced neutrophils enhanced metastasis after intravasation but before postextravasation colonization. Neutrophils protected cancer cells from natural killer (NK) cell–mediated clearance while still within the lumina of the microvessels. Overall, these results indicate that neutrophils can suppress the activity of NK cells and promote tumor cell extravasation. For details, please see the article by Spiegel and colleagues on page 630.

Cover image courtesy of Asaf Spiegel.