CAMB 7010: The Tumor Microenvironment

Directors: Ahmed Diab, PhD, Guilherme Nader, PhD, Ronny Drapkin, MD, PhD

Thursdays 3:30-5:30 pm Stellar Chance 204

Syllabus Spring 2025

Class Format:

- Students can meet with the assigned faculty prior to their presentation to discuss papers, slide deck, and any questions they may have.
- Students present background (30-40 minutes).
- 10-minute break
- Students present key data in paper(s) (45-50 minutes).
- Feedback from Ahmed, Guilherme, and Ronny to presenters only (10 minutes).

Each week, students that are not presenting that week will submit **two questions** they would like to discuss concerning impact/novelty/implications and/or general questions about the papers to Ahmed, Guilherme, and Ronny before or on the day of class.

If a student misses a class they are still expected to read the papers and submit their questions.

Course Grade: 40% assigned presentations, 40% overall class participation (and weekly questions), and 20% News and Views Article.

Classes:

(1/09/25) Class 1: Organizational Meeting

(1/23/25) Class 2: Inflammation and Tumor Progression (Guilherme)

Liudahl et al. "Leukocyte Heterogeneity in Pancreatic Ductal Adenocarcinoma: Phenotypic and Spatial Features Associated with Clinical Outcome", Cancer Discovery (2021).

Frittoli et al., "Tissue fluidification promotes a cGAS-STING cytosolic DNA response in invasive breast cancer" *Nature Materials* (2023).

(1/30/25) Class 3: Metastatic Niche (Ronny)

Lee et al. "Hepatocytes Direct the Formation of a Pro-metastatic Niche in the Liver", *Nature* (2019).

Fane et al. "Stromal Changes in the Aged Lung Induce an Emergence From Melanoma Dormancy", *Nature* (2022).

(2/06/25) Class 4: The Tumor Stroma, Cancer Associated Fibroblasts (Guilherme)

Barbazan et al., "Cancer-associated fibroblasts actively compress cancer cells and modulate mechanotransduction", *Nature Communications* (2023).

(2/13/25) Class 5: Crosstalk between the Tumor and the Microbiome (Ronny)

Yi et al. "Multimodal Immune Phenotyping Reveals Microbial-T cell Interactions That Shape Pancreatic Cancer", Cell Reports Medicine (2024).

"Targeting PD-L2-RGMb overcomes microbiome-related immunotherapy resistance", *Nature* (2023).

(2/20/25) Class 6: Immune Surveillance (Ahmed)

Herrera et al. "Low dose radiotherapy reverses tumor immune desertification and resistance to immunotherapy", Cancer Discovery (2021).

Li et al., "IGSF8 is an Innate Immune Checkpoint and Cancer Immunotherapy Target". *Cell* (2024).

(2/27/25) Class 7: Systemic Factors and Tumor Progression (Guilherme)

Argiles et al. "Cancer-associated cachexia — understanding the tumour macroenvironment and microenvironment to improve management", Nature Review Clinical Oncology (2023).

Queiroz et al. "Blocking ActRIIB signaling and restoring appetite reverses cachexia and improves survival in mice with lung cancer", *Nature Communications* (2022).

(3/06/25) Class 8: Tumor Angiogenesis, Lymphangiogensis (Ahmed)

Garcia Silva et al. "Melanoma-derived small extracellular vesicles induce lymphangiogenesis and metastasis through an NGFR-dependent mechanism", *Nature Cancer* (2021).

Stella Stasso "Lymphangiogenesis-inducing vaccines elicit potent and long-lasting T cell immunity against melanomas", *Science Advances* (2021).

--- Spring Break: March 8-16, 2025 ---

(3/20/25) Class 9: Stressful Tumor Microenvironments (Hypoxia and Nutrient Scarcity) (Ahmed)

Lien et al. "Low glycaemic diets alter lipid metabolism to influence tumour growth", *Nature* (2021).

Ubellacker et al. "Lymph protects metastasizing melanoma cells from ferroptosis", *Nature* (2020).

(3/27/25) Class 10: Tumor-Nervous System Interactions (Ronny)

Balood et al. "Nociceptor neurons affect cancer immunosurveillance", Nature 2022

Banh et al. "Neurons release serine to support mRNA translation in pancreatic cancer", *Cell* (2020).

(4/03/25) Class 11: The Influence of Age on Tumor Growth (Ronny)

Rossetti et al., "In Vivo DNA Replication Dynamics Unveil Aging-Dependent Replication Stress". *Cell* (2024).

Marino-Bravante et al. "Age-dependent Loss of HAPLN1 Erodes Vascular Integrity via Indirect Upregulation of Endothelial ICAM1 in Melanoma". *Nature Aging* (2024).

(4/10/25) Class 12: Tumor Metabolism (Ahmed)

Xiao et al. "Emerging therapies in cancer metabolism", Cell Metabolism (2023).

Tang et al. "Immunogenic coevolution defines unique microenvirnmental niches in ccRCC", Cell Metabolism (2023).

(4/17/25) Class 13: Tumor Dormancy (Guilherme)

Albrengues et al. "Neutrophil extracellular traps produced during inflammation awaken dormant cancer cells in mice", *Science* (2018).

Montagner et al., "Crosstalk with lung epithelial cells regulates Sfrp2-mediated latency in breast cancer dissemination", *Nature Cell Biology* (2020).

(4/24/25) Class 14: Cancer Heterogeneity, Plasticity, and Tumor Evolution (Guilherme)

Chia-Kuei Mo et al, "Tumour evolution and microenvironment interactions in 2D and 3D space", *Nature* (2024).

Concepcion et al. "SMARCA4 inactivation promotes lineage-specific transformation and early metastatic features in the lung," Cancer Discovery (2021).

(5/01/25) Class 15: Sex as a Biological Variable in Tumor Progression (Ahmed)

Vellano et al. "Androgen receptor blockade promotes response to BRAF/MEK-targeted therapy", *Nature* (2022).

Presentation Schedule:

Date	Background Presenter	Paper Presenter
1/23/25	Shirley Sun	Owen Koucky
1/30/25	Hira Khattak	Beatrice Goncalves
2/06/25	Grace Hinds	Madison Wahlsten
2/13/25	Jinfan Zhang	Shirley Sun
2/20/25	Mehrnaz Zakershahrak	Megan Costa
2/27/25	Bhargavi Subramanian	Jinfan Zhang
3/06/25	Owen Koucky	Grace Hinds
3/13/25	NO CLASS	NO CLASS
3/20/25	Beatrice Goncalves	Hira Khattak
3/27/25	Madison Wahlsten	Kim Manning
4/03/25	Megan Costa	Bhargavi Subramanian
4/10/25	Kim Manning	Mehrnaz Zakershahrak
4/17/25	Grace Hinds	Beatrice Goncalves
4/24/25	Kim Manning	Bhargavi Subramanian
5/01/25	Hira Khattak	Owen Koucky