



RESEARCH TOPIC MEM9
Role of epithelial plasticity in cancer initiation
Curriculum MEM

Laboratory name

Plasticity, Fibrosis and Cancer Lab Hunimed campus Bldg E, Pieve Emanuele

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Abstract

The Epithelial-to-Mesenchymal Transition (EMT) is an embryonic cellular trans-differentiation program relaunched in fibrosis and cancer. While its role in cancer progression, i.e. metastasis, is well established, little is known about its role during the early phases of neoplastic transformation. Recent evidences indicated that, rather than being a fibroblast-generating event, EMT represents an injury-induced response of the damaged epithelium impairing its functionality and regenerative capacity, and orchestrating inflammation. We seek to understand whether EMT activation could prime the injured epithelium to neoplastic transformation in those type of cancer arising from chronic and repeated inflammation such as colitis-associated colon cancer (CAC) in patients suffering from Inflammatory Bowel Disease (IBD).

The overall objective is to determine the functional consequences of the activation of the EMT program in terms of epithelial injury response, oncogenic priming and modulation of cancer-associated inflammation.

Main technical approaches

Cellular and molecular biology, use of in vivo preclinical models, histology, flow cytometry, confocal microscopy, image analysis

Scientific references

- 1) Lovisa S. Epithelial-to-Mesenchymal Transition in Fibrosis: Concepts and Targeting Strategies. *Front Pharmacol.* 2021 Sep 7;12:737570. PMID: 34557100
- 2) Lovisa S, Fletcher-Sananikone E, Sugimoto H, Hensel J, Lahiri S, Hertig A, Taduri G, Lawson E, Dewar R, Revuelta I, Kato N, Wu CJ, Bassett RL Jr, Putluri N, Zeisberg M, Zeisberg EM, LeBleu VS, Kalluri R. Endothelial-to-mesenchymal transition compromises vascular integrity to induce Myc-mediated metabolic reprogramming in kidney fibrosis. *Sci Signal.* 2020 Jun 9;13(635):eaaz2597. PMID: 32518142



3) Lovisa S, Genovese G, Danese S. Role of Epithelial-to-Mesenchymal Transition in Inflammatory Bowel Disease. *J Crohns Colitis*. 2019 Apr 26;13(5):659-668. PMID: 30520951.

4) Lovisa S, LeBleu VS, Tampe B, Sugimoto H, Vадnagara K, Carstens JL, Wu CC, Hagos Y, Burckhardt BC, Pentcheva-Hoang T, Nischal H, Allison JP, Zeisberg M, Kalluri R. Epithelial-to-mesenchymal transition induces cell cycle arrest and parenchymal damage in renal fibrosis. *Nat Med*. 2015 Sep;21(9):998-1009. PMID: 26236991

Type of contract

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