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Memorial Sloan Kettering Cancer Center

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Gabriela Chiosis, PhD

The goal of my program is to investigate stressor maladaptation mechanisms such as they occur in several diseases. Our approach takes advantage of the way nature has evolved to control such stressors, and that is by a unique usage of the chaperome, referred here as the epichaperome. The epichaperome, which we discovered, is structurally, dynamically and functionally distinct from the housekeeping chaperome, and my lab has pioneered an approach to take advantage of such features. By using innovative methods, we develop small molecule chemical toolsets specifically targeted to the epichaperome; these act as "sensors" of the epichaperome, and in turn, of the chronic stressor-associated proteome-wide malfunctions. By the use of these unique toolsets we aim to understand, diagnose and treat cellular processes associated with chronic stressors. We address multiple mechanistic and biochemical questions less amenable to approaches that treat the chaperome as monolithic entity (*i.e.* the classical biochemical and genetic tools). We investigate in endogenous systems, both at the cellular and the

organismal level, the inherent proteome changes and mechanisms that lead to disease, *i.e.* we can understand. By sensing disease states through the chemical toolsets, we go beyond investigation; we identify, measure and quantify, *i.e.* we can diagnose. By attacking the epichaperome specifically, we perturb the disease-causing proteome, and in turn revert or slow the disease phenotype, *i.e.* we can treat.

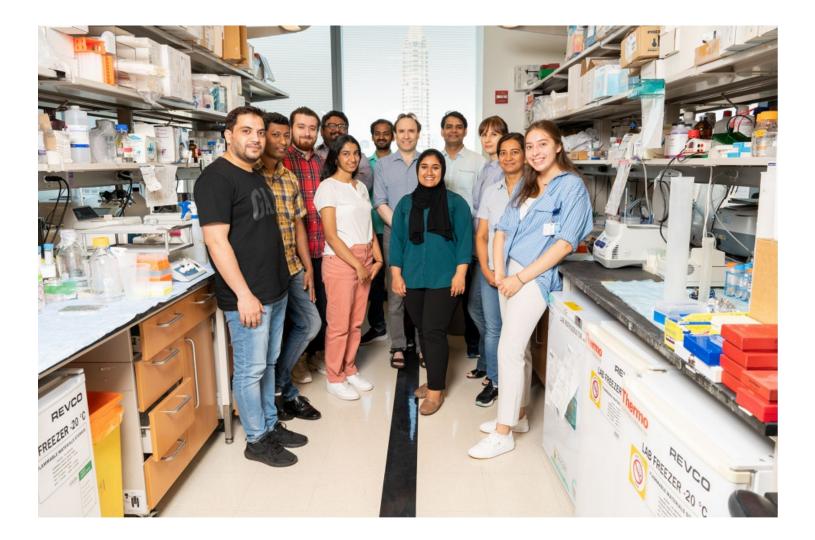
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Research Projects

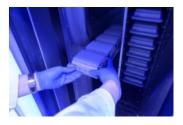
Protein-protein interaction networks in disease Mechanisms of disease Epichaperomes in Neurodegenerative Diseases Paradigms for Precision Medicine Development of Chemical Probes

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Featured News



MSK Research Highlights, June 29, 2023

New MSK research discovered ferroptosis regulators that suggest therapeutic opportunities against hormone receptor-positive cancers; examined how tumor-associated macrophages might be turned against cancer; acquired new insights into joint inflammation in rheumatoid arthritis; developed a systems-level platform called epichaperomics to map changes in interactors among thousands of proteins involved in cancer-related processes; and investigated how artificial intelligence could help diagnose an invasive form of breast cancer.

IN THE LAB



Experimental Drug Targets Misbehaving Proteins in Brain Cancer and Alzheimer's Disease

Memorial Sloan Kettering researchers are studying how drugs that reverse malfunctioning proteins may treat disease.

IN THE LAB



Just Add Sugar: How a Protein's Small Change Leads to Big Trouble for Cells

A study from investigators in the Sloan Kettering Institute uncovers the details of how a key protein called GRP94 becomes disrupted, leading to cancer and other diseases.

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Publications Highlights

The epichaperome is an integrated chaperome network that facilitates tumour survival. Rodina A, Wang T, Yan P, Gomes

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People

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The Chiosis lab uses a unique chemical biology approach to understand, diagnose, and treat cellular processes associated with chronic molecular stress, with the ultimate goal of developing novel therapeutic options for use in the clinic. PhD, Columbia University

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Palak Panchal Research Technician



Julia Ashmead Associate Administrative Assistant

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AACR – Cancer Research and Prevention Career Development Award in Translational Lung Cancer Research, in Memory of Duffy Wall

Susan G. Komen Breast Cancer Translational Research Award

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Top 5 percent cited author in Biology and Biochemistry 2010 (analysis by Thomson Reuters)

Translated from bench-to-bedside the Hsp90 inhibitor PU-H71 and the non-invasive companion diagnostic 124I-PU-H71 PET

Lab News & Events



A new strategy for Alzheimer's disease treatment targets cell-wide protein malfunction

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Gabriela Chiosis discloses the following relationships and financial interests:

Samus Therapeutics LLC Equity; Intellectual Property Rights

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