

# MIT Industrial Liaison Program Faculty Knowledgebase Report

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2022 MIT Health Science Technologies Conference

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April 12, 2022 9:00 am - 5:00 pm

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8:00 AM - 9:00 AM

Registration with Light Breakfast

Welcome and Introduction  
John Roberts  
Executive Director (Interim), [MIT Corporate Relations](#)



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Executive Director (Interim)  
[MIT Corporate Relations](#)

John Roberts has been Executive Director of MIT Corporate Relations (Interim) since February 2022. He obtained his Ph.D. in organic chemistry at MIT and returned to the university after a 20-year career in the pharmaceutical industry, joining the MIT Industrial Liaison Program (ILP) in 2013. Prior to his return, John worked at small, medium, and large companies, holding positions that allowed him to exploit his passions in synthetic chemistry, project leadership, and alliance management while growing his responsibilities for managing others, ultimately as a department head. As a program director at MIT, John built a portfolio of ILP member companies, mostly in the pharmaceutical industry and headquartered in Japan, connecting them to engagement opportunities in the MIT community. Soon after returning to MIT, John began to lead a group of program directors with a combined portfolio of 60-80 global companies. In his current role, John oversees MIT Corporate Relations which houses ILP and MIT Startup Exchange.

Sheryl Greenberg  
Program Director, [MIT Industrial Liaison Program](#)



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Program Director  
[MIT Industrial Liaison Program](#)

Sheryl Greenberg initiates and promotes the interactions and development of relationships between academic and industrial entities to facilitate the transfer of new ideas and technologies between MIT and companies, and has created numerous successful partnerships. By understanding the business, technology, and commercial problems within a company, and understanding the technologies and expertise of MIT researchers, Greenberg identifies appropriate resources and expertise to foster new technology applications and collaborative opportunities.

Prior to MIT, Greenberg created and directed the Office of Technology Transfer at Brandeis University. In the process of managing intellectual property protection, marketing, and licensing, she has promoted the successful commercialization of technologies as diverse as new chemicals and manufacturing, biotechnology, food compositions, software, and medical devices. She facilitated the founding and funding of new companies, as well as creating a profitable technology transfer program. She also facilitated the patenting, marketing, and licensing of Massachusetts General Hospital technologies. In addition to her cellular, biochemical, and genetic research experience in academic and corporate environments, she has also created intellectual property for medical uses. Greenberg has been an independent intellectual property and business development consultant, is a U.S. Patent Agent, and has previously served the Juvenile Diabetes Research Foundation as Co-Chair of the Islet Research Program Advisory Committee and grant reviewer. She currently also mentors startup companies and facilitates partnering them with large life science and healthcare companies.

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9:20 AM - 9:50 AM	<p>DFCI: The Challenges and Opportunities in Cell and Gene Therapy</p> <p>Eric Smith          Director of Translational Research, Immune Effector Cell Therapies?          Principal Investigator of a Synthetic Biology/Cellular Engineering/mRNA laboratory  <a href="#">Dana-Farber Cancer Institute</a></p>
	<p>CAR T cell therapies for relapsed/refractory hematologic malignancies induce frequent deep clinical responses in patients who failed all other treatment options. While these results are game changing, the field is only scratching the surface of the potential benefit gene and cell therapies can provide. Future advances include discovery of novel target antigens, extension of CAR T cell clinical benefit to solid tumors, logic gated engineering designs, advances in manufacturing, targeted in situ gene delivery, and off-the-shelf immunologically stealth approaches. The above will be broadly addressed and work from our group developing and translating CAR T cell therapies targeting the novel antigen GPRC5D for myeloma and "OR" gating strategies for hematologic malignancies will be highlighted as examples.</p>
9:50 AM - 10:20 AM	<p>Synthetic Gene Networks for Multi-step Differentiation of Stem Cells into Programmable Organoids</p> <p>Deepak Mishra          Instructor and Postdoctoral Associate in Biological Engineering, MIT</p>
	<p>Organoids are tiny, self-organized three-dimensional (3D) tissue cultures derived from stem cells that allow recapitulation of architecture, composition, and physiology of organs. Organoids may be formed by combining individually differentiated cell populations or via the introduction of external cues to human induced pluripotent stem cells (hiPSCs) to mimic stages of human development. However, such approaches may limit intercellular interactions or faithful mimicry of a mature organ's structure. We employ synthetic gene circuits to genetically engineer human induced pluripotent stem cells (hiPSCs) with transcriptional and post-transcriptional regulatory elements. These networks govern multi-step differentiation of hiPSCs in a cell-specific, semi-automated manner to form 3D liver organoids. Our recent advances in creation and subsequent use of these organoids in toxicology, drug metabolism, and implantation will be presented and discussed.</p>
10:20 AM - 10:50 AM	<p>Self-Replicating RNA Technologies for Vaccines and Cancer Immunotherapy</p> <p>Parisa Yousefpour          Postdoctoral Associate, Darrell Irvine Lab  <a href="#">Koch Institute for Integrative Cancer Research at MIT</a></p>
	<p>Self-replicating RNAs termed replicons have begun to be explored as a promising platform technology for vaccines and gene therapy. Upon delivery to host cells, the replicon copies itself and therefore, allows for prolonged and increased transgene expression with a small initial dose. We employ the replicon platform for 1) vaccine development for sustained antigen expression coupled with the intrinsic adjuvanticity of replicons, and 2) cancer immunotherapy to stimulate multiple synergistic pathways of antitumor immunity. In addition, harnessing synthetic biology tools, we are developing next generation replicon platforms that incorporate microRNA-based classifier and small-molecule responsive gene circuits for internal and external regulation of transgene expression, respectively. Our recent advances and directions on gene delivery with replicons will be presented and discussed.</p>
10:50 AM - 11:10 AM	<p>Networking Break</p>
11:10 AM - 11:15 AM	<p>MIT Professional Education</p> <p>Myriam Joseph          Assistant Director, Marketing &amp; Business Development  <a href="#">MIT Professional Education</a></p>

11:15 AM - 12:15 PM

MIT Startup Exchange Lightning Talks  
Ariadna Rodenstein  
Program Manager, [MIT Startup Exchange](#)



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Program Manager  
[MIT Startup Exchange](#)

Ariadna Rodenstein is a Program Manager at MIT Startup Exchange. She joined MIT Corporate Relations as an Events Leader in September 2019 and is responsible for designing and executing startup events, including content development, coaching and hosting, and logistics. Ms. Rodenstein works closely with the Industrial Liaison Program (ILP) in promoting collaboration and partnerships between MIT-connected startups and industry, as well as with other areas around the MIT innovation ecosystem and beyond.

Prior to working for MIT Corporate Relations, she worked for over a decade at Credit Suisse Group in New York and London, in a few different roles in event management and as Director of Client Strategy. Ms. Rodenstein has combined her experience in the private sector with work at non-profits as a Consultant and Development Director at New York Immigration Coalition, Immigrant Defense Project, and Americas Society/Council of the Americas. She also served as an Officer on the Board of Directors of the Riverside Clay Tennis Association in New York for several years. Additionally, she earned her B.A. in Political Science and Communications from New York University, with coursework at the Instituto Tecnológico y de Estudios Superiores de Monterrey in Mexico City, and her M.A. in Sociology from the City University of New York.

Paulo Garcia  
CEO & Co-founder  
[Kytopen](#)

Marinna Madrid  
Co-founder  
[Cellino](#)

Gopi Shanker  
Chief Scientific Officer  
[Tevard Biosciences](#)

Fred Parietti  
Founder & CEO  
[Multiply Labs](#)

Floris Engelhardt  
CEO  
[Kano Therapeutics](#)

Udayan Umapathi  
Co-founder & CEO  
[Volta Labs](#)

Daniel Meyer  
CEO  
[CellChorus](#)

Elisabeth Maida  
Co-founder & CEO  
[Fathom Data](#)

12:15 PM - 1:25 PM

Lunch with Startups Exhibit

**Exhibit only startup**

[LiquiGlide](#): Friction-reducing coating technology to enhance cell/gene delivery performance

1:30 PM - 2:00 PM

### Unlocking the Potential of Life Sciences Innovation by Harnessing the Power of Mission-driven Collaborations

Ran Zheng  
CEO  
[Landmark Bio](#)

We have entered the golden age of biology with an unprecedented explosion of discoveries that could lead to life changing therapies and have profound impact on human health. However, the shortage of manufacturing capacity and expertise in recent years has constrained therapeutic translation especially for the life sciences innovators who explore emerging modalities and unique technologies. Manufacturing scalability, reproducibility and the cost of goods remain as major challenges, while the evolving science and regulatory landscape require continuous learning.

In 2021, Harvard and MIT partnered with Cytiva, Fujifilm Diosynth Biotechnologies and Alexandria Real Estate Equities to bring the best of academia, the life sciences industry, and world-class research hospitals together to accelerate life sciences innovation. Landmark Bio was established to break down the barriers in novel therapeutics development and industrialization. We collaborate with academics, startups and drug developers to take groundbreaking discoveries from bench to clinics by providing capacity and expertise in CMC development and manufacturing. Uniquely positioned as pre-platform and technology agnostic, we help design, de-risk, and develop early innovations into future platforms. Our cross-sector partnership offers an unparalleled academic, industry, hospitals and investor network with a magnetic pull to power biomanufacturing innovation. As a mission-driven organization, we embrace risk and opportunities at the bleeding-edge in order to advance emerging technologies, demonstrate therapeutic potential and improve human health.

2:00 PM - 2:30 PM

### Label-Free Biophysical Critical Quality Attributes (CQAs) for Cell Therapy Products Jongyoon Han

Professor of Electrical Engineering, [Department of Electrical Engineering & Computer Science \(EECS\)](#)  
Professor of Biological Engineering, [Biological Engineering Division](#)



Jongyoon Han

Professor of Electrical Engineering, [Department of Electrical Engineering & Computer Science \(EECS\)](#)  
Professor of Biological Engineering, [Biological Engineering Division](#)

Dr. Jongyoon Han is currently a professor in the Department of Electrical Engineering and Computer Science and the Department of Biological Engineering, Massachusetts Institute of Technology. He received B.S.(1992) and M.S.(1994) degree in physics from Seoul National University, Seoul, Korea, and Ph.D. degree in applied physics from Cornell University in 2001. He was a research scientist in Sandia National Laboratories (Livermore, CA), until he joined the MIT faculty in 2002. He received NSF CAREER award (2003) and Analytical Chemistry Young Innovator Award (ACS, 2009). His research is mainly focused on applying micro/nanofabrication techniques to a very diverse set of fields and industries, including biosensing, desalination / water purification, biomanufacturing, dentistry, and neuroscience. He is currently the lead PI for MIT's participation for NIIMBL (The National Institute for Innovation in Manufacturing Biopharmaceuticals).

One of the critical challenges in cell therapy is the lack of reliable, specific, and non-destructive quality attributes, which are sorely needed for all aspects of biomanufacturing of these cells, including donor selection, in-process quality monitoring, and release testing. Most biological and biochemical assays are destructive or perturbative, which limits their utility, especially for autologous cell therapy. In this talk, I will showcase some of the emerging ideas of label-free, biophysical critical quality attributes (CQAs) we have been working on, including magnetic, electrical, and mechanical signatures of cells. Once the strong correlation with biochemical cell phenotypes, these will serve an important role to improve the overall production of both allogeneic and autologous cell therapy products.

2:30 PM - 3:00 PM

Exploration of Biological Diversity

Feng Zhang

Core Member, Broad Institute of MIT and Harvard

Investigator, McGovern Institute for Brain Research, MIT

James and Patricia Poitras Professor of Neuroscience at MIT

Associate Professor, Departments of Brain and Cognitive Sciences and Biological Engineering, MIT

New York Stem Cell Foundation-Robertson Investigator



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Associate Professor, Departments of Brain and Cognitive Sciences and Biological Engineering, MIT

New York Stem Cell Foundation-Robertson Investigator

Feng Zhang attended Harvard University as an undergraduate, where he earned an A.B. in Chemistry and Physics in 2004. He transitioned to Stanford for his Ph.D. in Chemistry and Biophysics, which he obtained in 2009 while working in Karl Deisseroth's research group for his work pioneering optogenetics in conjunction with Ed Boyden. After a postdoc in George Church's lab, he moved to MIT in January of 2011, where he is now both an Associate Professor of Biological Engineering and a Core Institute Member of the Broad Institute of MIT and Harvard.

Feng Zhang has received a number of honors, including the NIH Pioneer Award and the Perl-UNC Neuroscience Prize, along with Ed Boyden and thesis advisor Deisseroth. He is additionally a Searle Scholar.

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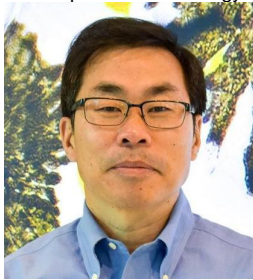
Many powerful molecular biology tools have their origin in nature, and, often, microbial life. From restriction enzymes to CRISPR-Cas9, microbes utilize a diverse array of systems to get ahead evolutionarily. We are interested in exploring this natural diversity through bioinformatics, biochemical, and molecular work to better understand the fundamental ways in which living organisms sense and respond to their environment and ultimately to harness these systems to improve human health. Building on our demonstration that Cas9 can be repurposed for precision genome editing in mammalian cells, we began looking for novel CRISPR-Cas systems that may have other useful properties. This led to the discovery of several new CRISPR systems, including the CRISPR-Cas13 family that target RNA, rather than DNA. We developed a toolbox for RNA modulation based on Cas13, including methods for precision base editing. We are expanding our biodiscovery efforts to search for new microbial proteins that may be adapted for applications beyond genome and transcriptome modulation, capitalizing on the growing volume of microbial genomic sequences and building on our bioengineering expertise. We are particularly interested in identifying new therapeutic modalities and vehicles for delivering cellular and molecular cargo. We hope that this combination of tools and delivery modes will accelerate basic research into human disease and open up new therapeutic possibilities.

3:00 PM - 3:20 PM

Networking Break

CAR-NK Cell Therapies for Cancer

Jianzhu Chen  
Professor of Biology, Koch Institute for Integrative Cancer Research  
MIT Department of Biology



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Professor of Biology, Koch Institute for Integrative Cancer Research  
MIT Department of Biology

Jianzhu Chen is Professor of Biology at Koch Institute for Integrative Cancer Research and Department of Biology at Massachusetts Institute of Technology (MIT). Dr. Chen received a Ph.D. degree from Stanford University. He was a postdoctoral fellow and then an instructor at Harvard Medical School before he joined the faculty at MIT. Dr. Chen's research seeks fundamental understanding of the immune system as well as its application in disease intervention. Over the years, Dr. Chen has made significant contributions to a broad area of research in immunology, cancer research, infectious diseases, and animal models of human diseases. His research on lymphocyte homeostasis and immunological memory challenged the prevailing paradigm at the time and revealed unexpected effect of lymphopenia-induced proliferation on memory T cell development. Dr. Chen pioneered innovative research in development of novel mouse models through genetic manipulations and cell complementation and in development of novel anti-microbial, including surface coatings that inactivate microbes on contact, universal anti-influenza siRNAs, and polymer-attached antivirals that minimize drug resistance of influenza viruses. Recently, Dr. Chen's research activity has focused on development of tumor-specific CAR-NK cells and re-programming macrophages for disease intervention, including cancer, metabolic diseases and infectious diseases.

[View full bio](#)

Natural Killer (NK) cells and CD8+ cytotoxic T cells are two types of immune cells that can kill target cells through similar cytotoxic mechanisms. With the remarkable success of chimeric antigen receptor-engineered T (CAR-T) cells for treating hematological malignancies, there is a rapidly growing interest in developing CAR-engineered NK (CAR-NK) cells for cancer therapy. Compared to CAR-T cells, CAR-NK cells could offer some significant advantages, including (1) better safety, such as a lack of or minimal cytokine release syndrome and neurotoxicity in autologous setting and graft-versus-host disease in allogeneic setting, (2) multiple mechanisms for activating cytotoxic activity, and (3) high feasibility for "off-the-shelf" manufacturing. We are developing the next generation of CAR-NK cells by combining tumor-specific CAR, additional armors, and cytokine-induced memory-like (CIML) NK cells, with a goal to achieve better tumor-specific targeting, enhanced proliferation and persistence *in vivo*, resistance to the suppressive tumor microenvironment, and ultimately an effective and durable anti-tumor response in patients.

Rational Design of rAAV Production via Mechanistic Modeling

Richard Braatz

Edwin R. Gilliland Professor, [MIT Department of Chemical Engineering](#)



Richard Braatz

Edwin R. Gilliland Professor

[MIT Department of Chemical Engineering](#)

Dr. Richard D. Braatz is the Edwin R. Gilliland Professor of Chemical Engineering at MIT, where he conducts research into advanced biomanufacturing systems. He is the Director of the Center on Continuous mRNA Manufacturing and leads process data analytics, mechanistic modeling, and control systems for projects on vaccine, monoclonal antibody, and gene therapy manufacturing. Dr. Braatz received an M.S. and Ph.D. from the California Institute of Technology and was the Millennium Chair and Professor at the University of Illinois at Urbana-Champaign and a Visiting Scholar at Harvard University before moving to MIT. Dr. Braatz has collaborated with more than 20 companies, including Novartis, Pfizer, Merck, Bristol-Myers Squibb, Biogen, Amgen, Takeda, and Abbott Labs. He has published over 300 papers and three books. Dr. Braatz is a Fellow of IEEE, IFAC, AIChE, and AAAS and a member of the U.S. National Academy of Engineering.

[View full bio](#)

Tam Nguyen

Ph.D. student in chemical engineering at MIT

Recombinant adeno-associated virus (rAAV) is one of the most commonly used platforms for *in vivo* gene therapy treatments. The reduced toxicity, robust and long-term transgene expression, and ability to transduce both dividing and non-dividing cells as well as target a wide range of tissues have made rAAV the most widely used viral vector. However, the standard method of producing rAAV via transient transfection of mammalian cells, specifically human embryonic kidney 293 (HEK293) cells, typically has low yield and generates a high portion of empty particles, laying extra burden on downstream processing. To elucidate the mechanisms of rAAV synthesis in HEK293 suspension-adapted cells, we have developed a mechanistic model based on the published understanding of the underlying biology and existing data. Quantitative analysis suggests the misaligned dynamics of capsid and viral DNA production result in the high ratio of empty particles. Through a model-based strategy, we explored a novel transfection method using low-dose multiple transfections in HEK293 cell culture that successfully increased the ratio of full to empty capsids in the viral harvest without compromising the viral titer. Molecular analysis through a next-generation rAAV production model attributed the improvements to changes in the kinetics of viral protein expression and DNA replication. Here, we demonstrate that the use of multiple transfection times is a practical method for increasing the genome titer and improving the percentage of full capsids for rAAV production. Our results also demonstrated the capability to manipulate product composition from an operational standpoint.



4:20 PM - 4:50 PM

## Reprogramming Adaptive Immunity

Michael Birnbaum

Associate Professor of Biological Engineering, [MIT Department of Biological Engineering](#)



Michael Birnbaum

Associate Professor of Biological Engineering

[MIT Department of Biological Engineering](#)

Michael obtained an A.B. in Chemical and Physical Biology at Harvard University in 2008. He then moved to Stanford University, where he completed his Ph.D. in Immunology in 2014. At Stanford, he worked in Professor K. Christopher Garcia's laboratory, studying the molecular mechanisms of T cell receptor recognition, cross-reactivity, and activation. He then conducted postdoctoral research in Professor Carla Shatz's laboratory, studying novel roles for immune receptors expressed by neurons in neural development and neurodegenerative disease. Michael joined the Department of Biological Engineering in 2016 as an Assistant Professor.

[View full bio](#)

Cell and gene immunotherapies are revolutionizing how we treat disease, with multiple FDA-approved therapies that have transformed cancer treatments. However, advances in gene delivery, manufacturing, and therapeutic cargoes are still required to increase the impact and scope of these promising approaches. The Birnbaum laboratory is working to develop approaches that improve the specificity of cellular engineering, and the potency of cells once engineered.

4:50 PM - 6:00 PM

Closing Remarks and Networking Reception