

Viral Vector Free Cellular Engineering

Bruce L. Levine, Ph.D.

Center for Cellular Immunotherapies, University of Pennsylvania

Co-Founder, Tmunity Therapeutics

President, International Society for Cell and Gene Therapy



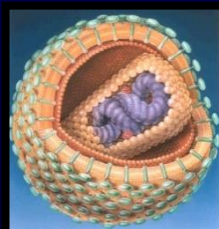
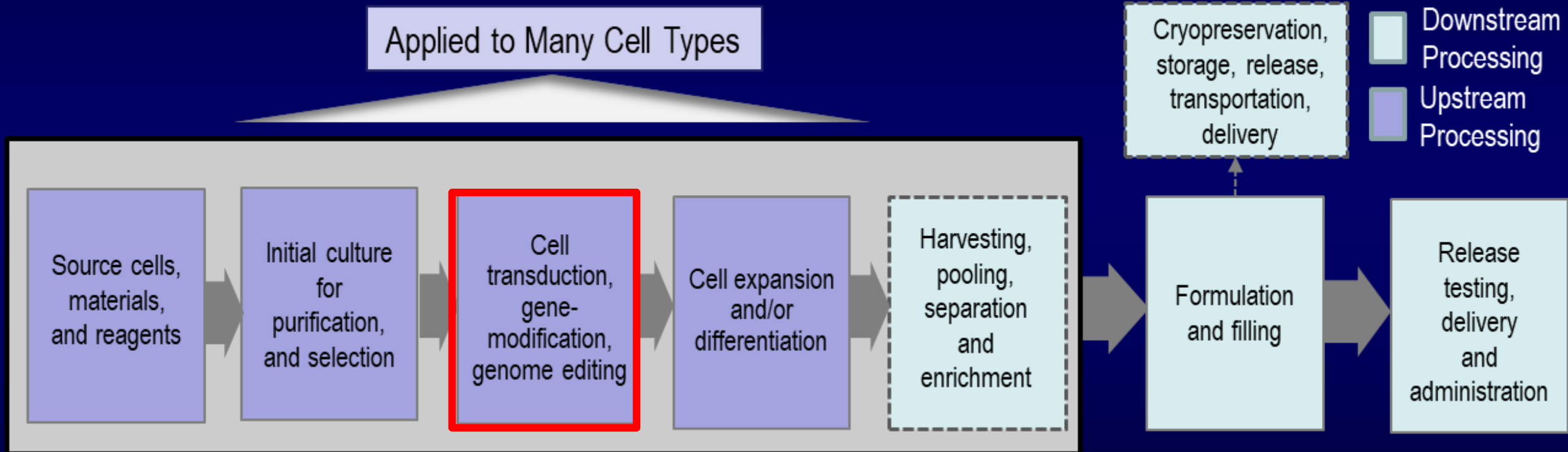
@BLLPHD



Conflict of Interest Statement

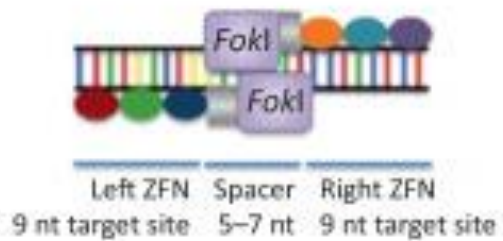
- Declaration of financial interest due to intellectual property and patents in the field of cell and gene therapy.
- University of Pennsylvania Alliance with Novartis
- Consultant for Terumo
- Scientific Advisory Board for Akron, Avectas, Immuneel, Immusoft, In8bio, Ori Biotech, Vycellix
- Co-Founder and equity holder Tmunity Therapeutics
- Conflict of interest is managed in accordance with University of Pennsylvania policy and oversight

Engineered Immune Cell Ex Vivo Processing Flow

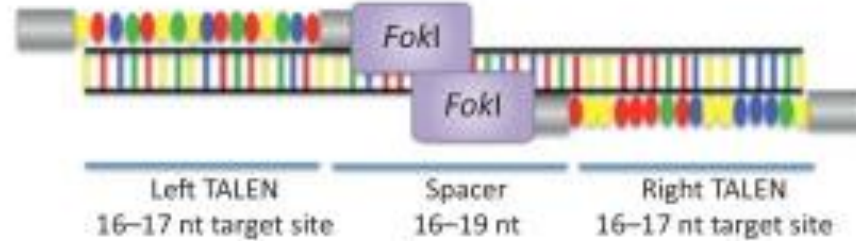


Genomic disruption (and insertion) as therapeutic cell engineering tools

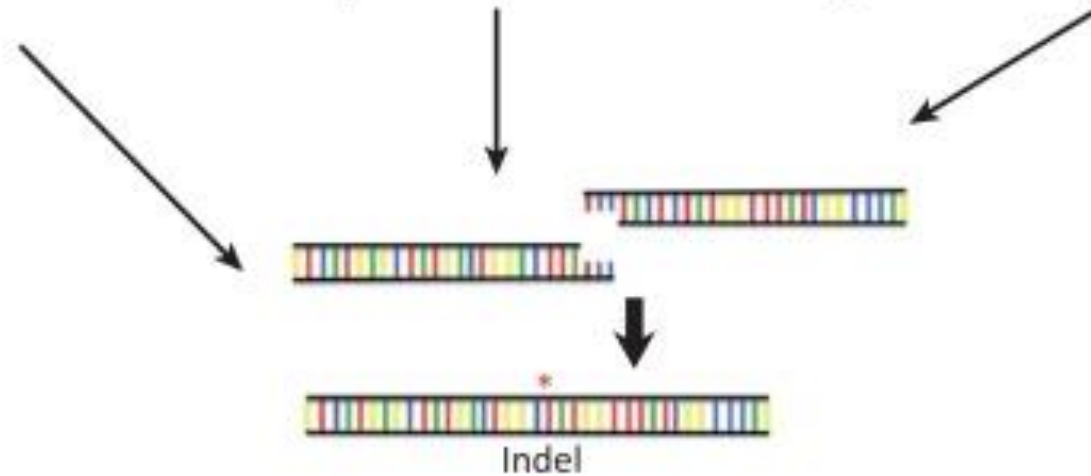
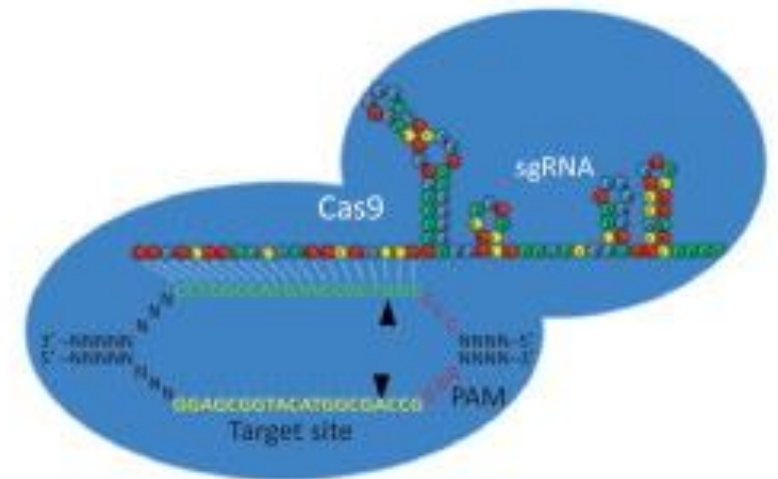
(A) ZFN
Zinc finger nuclease



(B) TALEN
Transcription activator-like effector nuclease

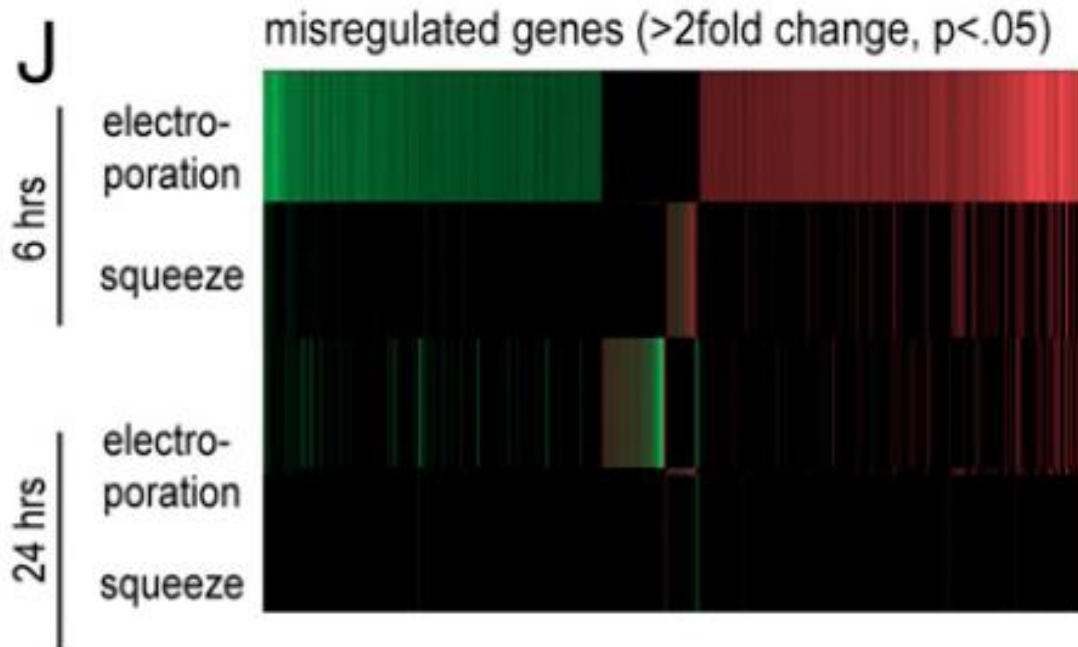
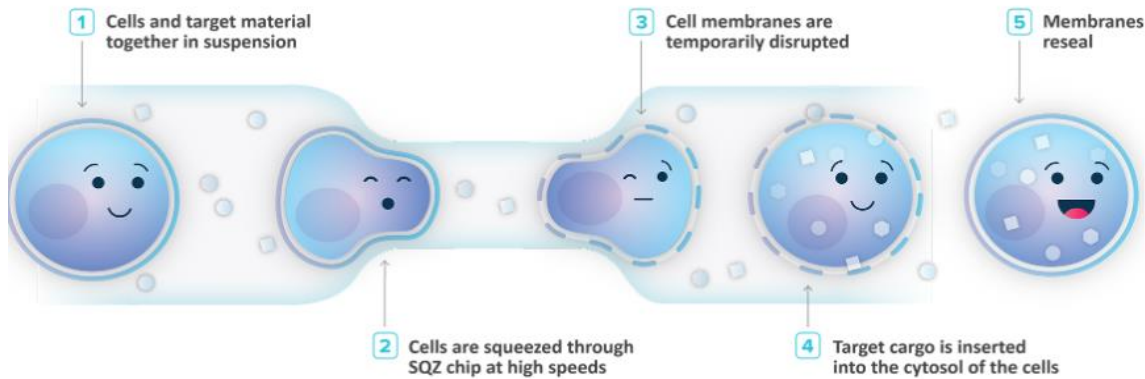


(C) CRISPR-Cas system

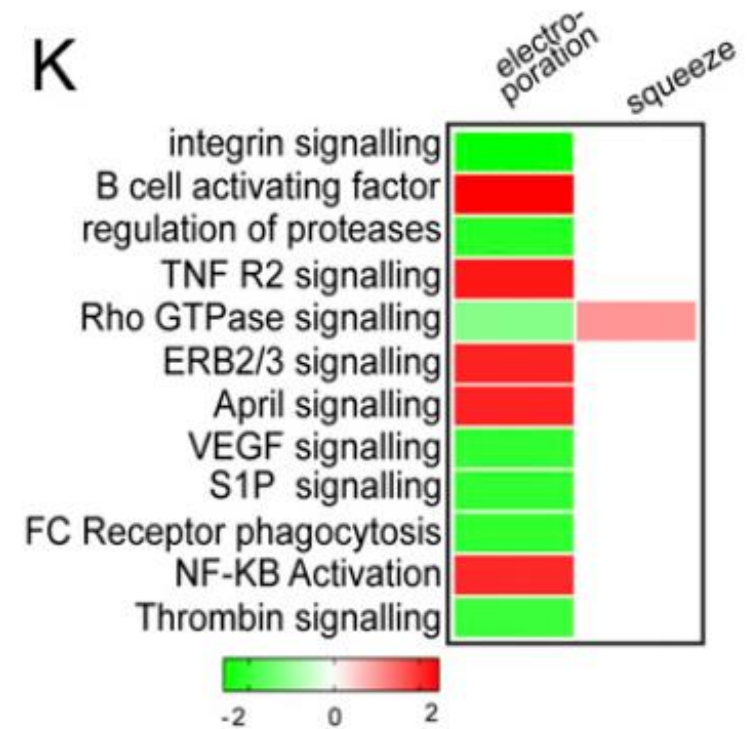


Mechanical Poration

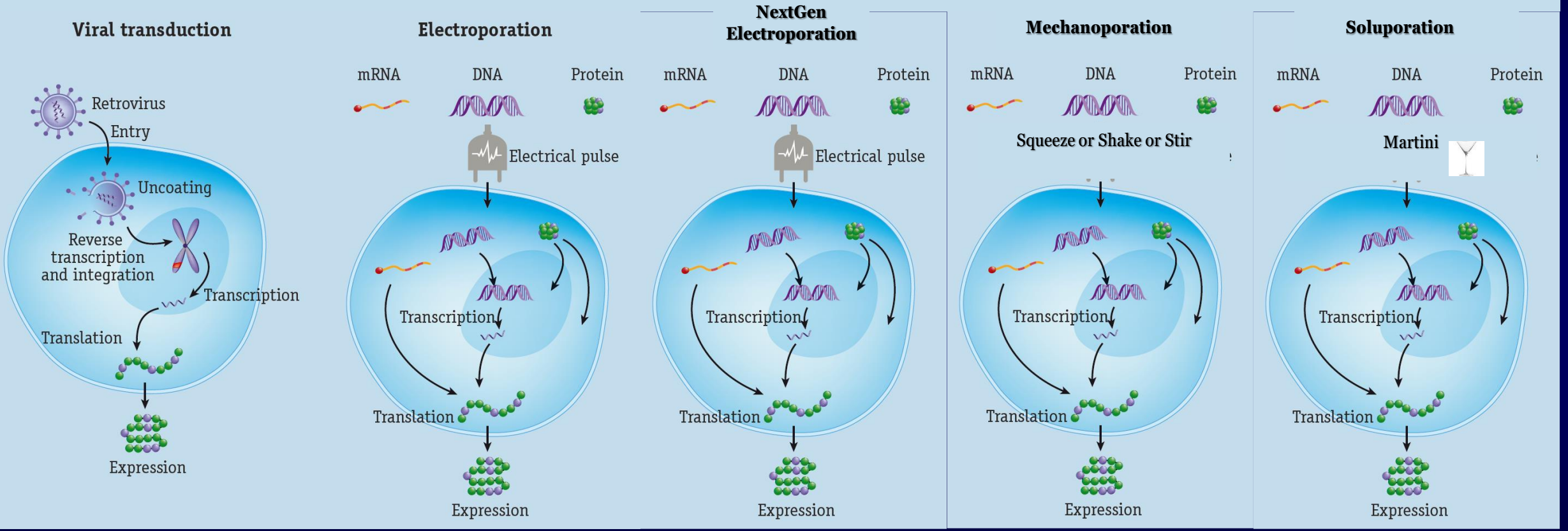
SQZ, MIT spinout



K



Cellular Door Dash: Gene/Cargo Delivery

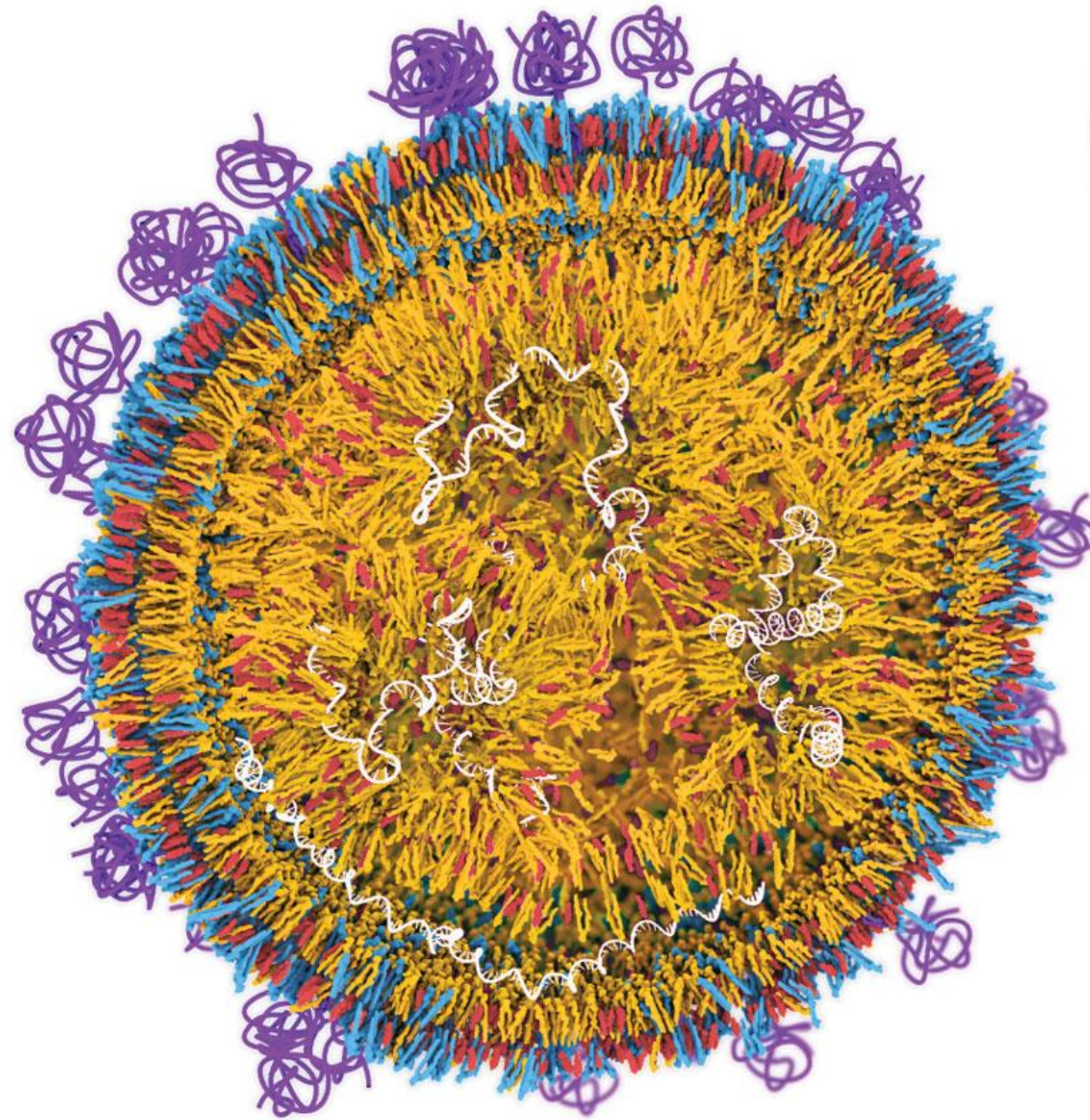


**MIT
Kytopen**

**Georgia Tech
CellFE**

**Maynooth Univ.
Avectas**

Vector free technology of the year/decade



Lipid nanoparticle, 80 nm wide



**2-6
mRNA strands**



**1,000
PEG lipids**



**6,000
Phospholipids**



**20,000
Cholesterol molecules**



**30,000
Ionizable lipids**

Electro-Mechanical Transfection

Cullen R. Buie, PhD | Massachusetts Institute of
Technology

Transient Cell Volume Exchange

Todd Sulchek, PhD | Georgia Institute of Technology

Enabling the Next Wave of Non-Viral Cell Therapies

Michael Maguire, PhD, CEO | Avectas