

A DNA-Encoded Library (DEL) Platform for Academic Targets

Alexander L. Satz May 26, 2020

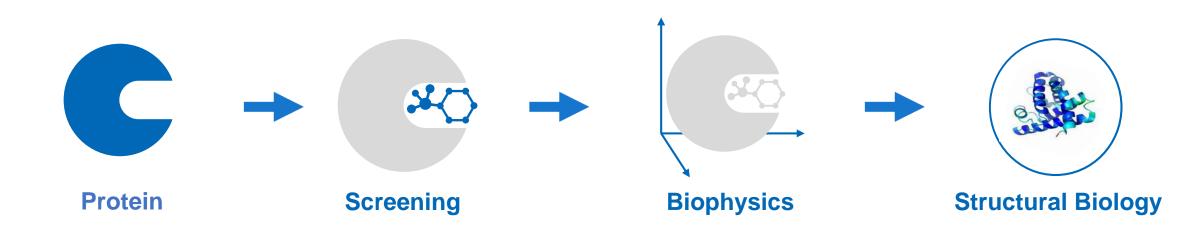




Targets to Chemical Matter

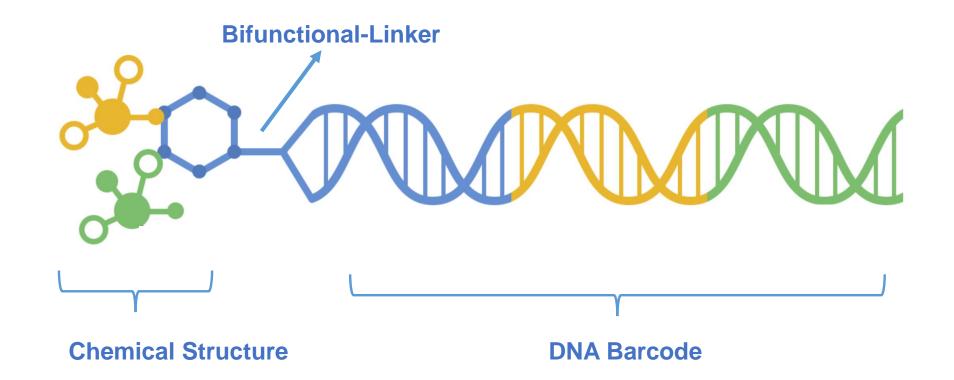
Early Stage Drug Discovery





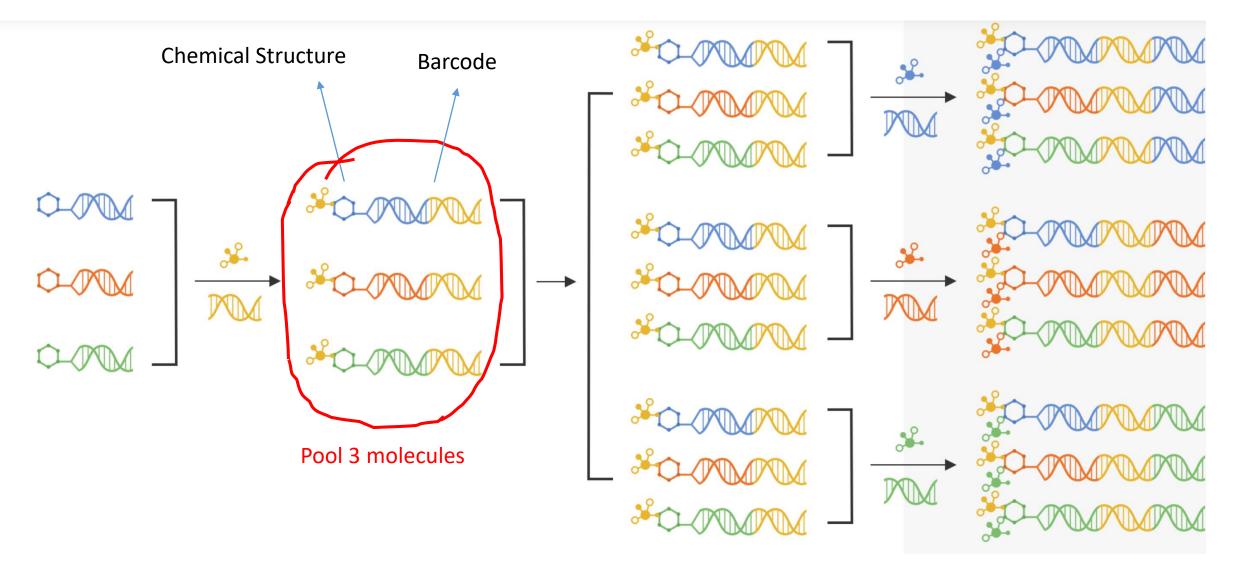
DNA Encoded Libraries (DEL) Molecule



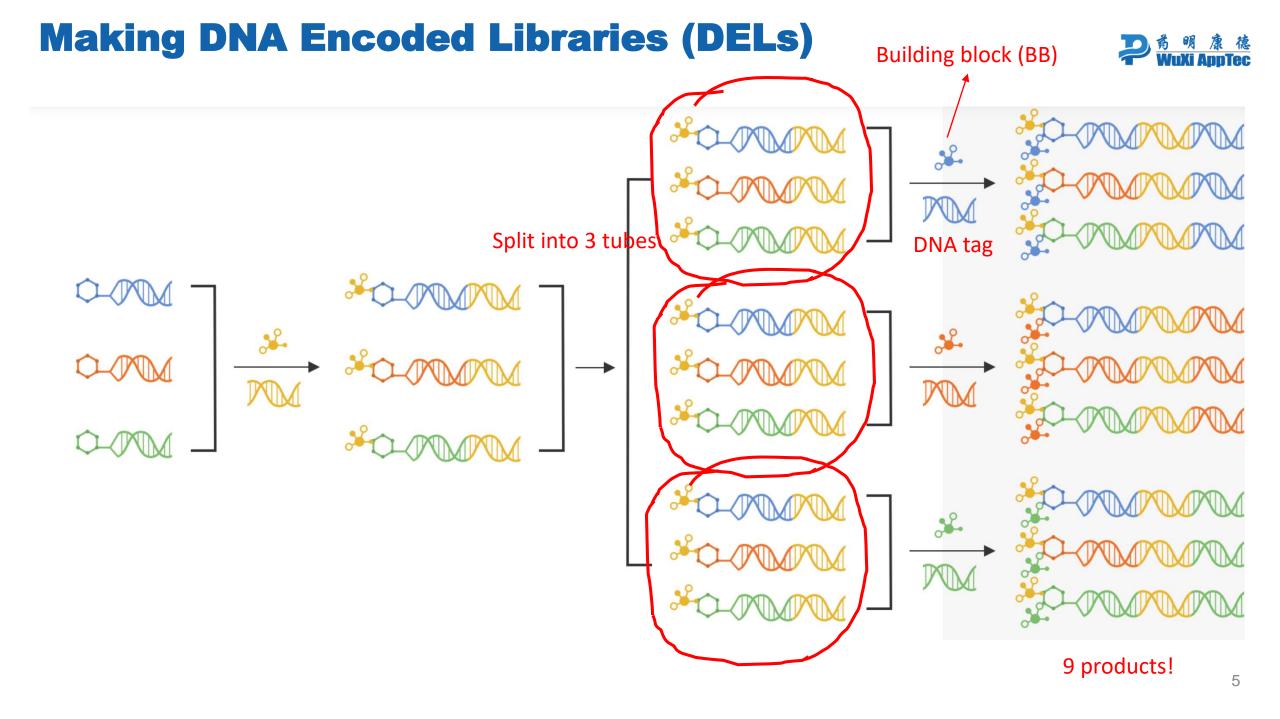


Making DNA Encoded Libraries (DELs)

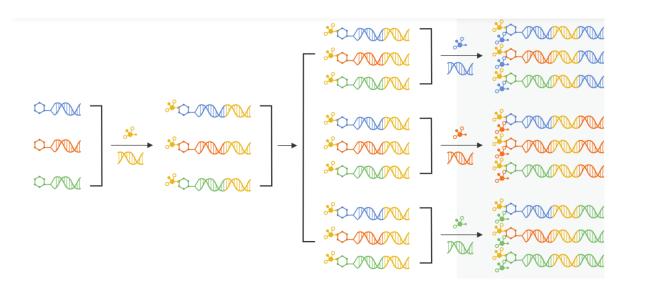




'Split-and-Pool Combinatorial Chemistry'



Making DNA Encoded Libraries (DELs)



96 x 96 x 96 x 96 = 85 million different chemical structures

Or

$9000 \times 9000 = 81$ million structures

Billions of chemical structures in each tube

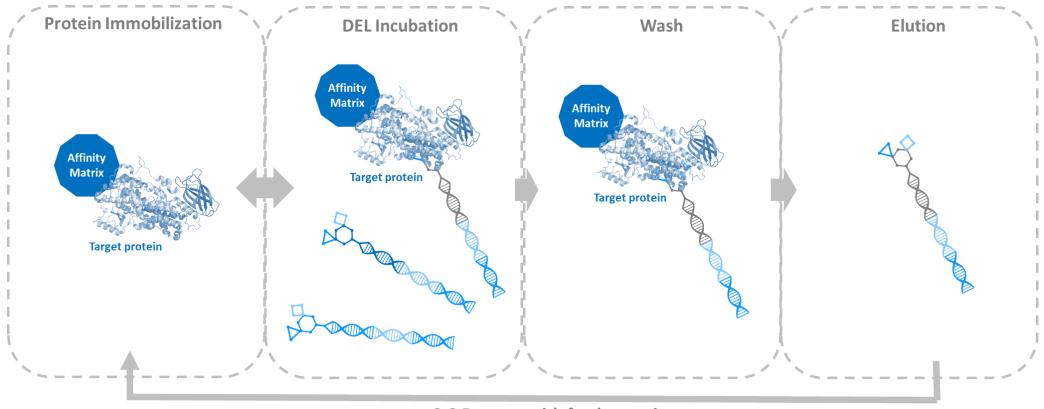




14 billion different encoded chemical structures in each tube!

Each tube contains (effectively) an identical copy of the same 14 billion chemical structures

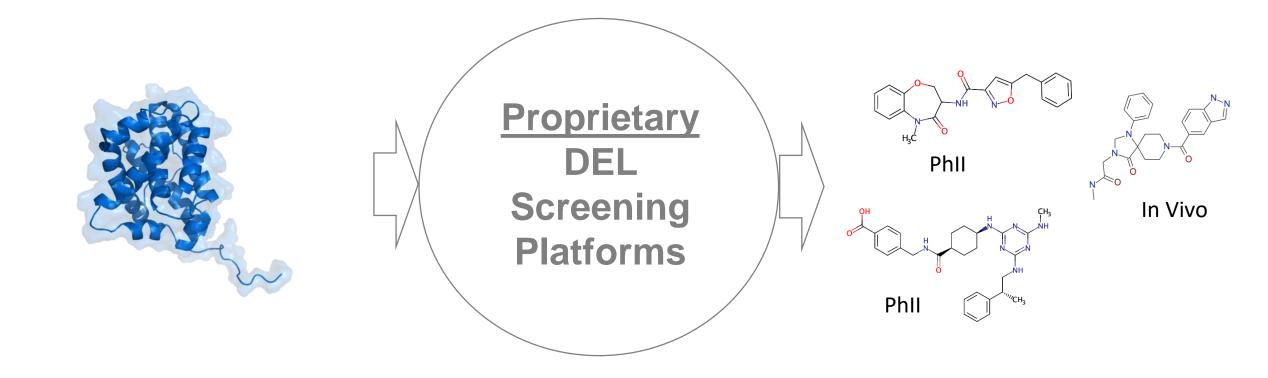
DEL Affinity Screen



2-3 Repeats with fresh protein

DEL Screening Provides Drug-like Chemical Matter





For More Examples of Published DEL Hits:

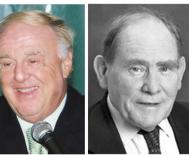
What Do You Get from DNA-Encoded Libraries? Satz, A. ACS Med. Chem. Lett., 2018; 9(5), 408-410. Chemical Space of DNA-Encoded Libraries. Franzini, R.M. and Randolph, C., J Med Chem, 2016; 59(14), 6629-44.

Open Access to DEL Screening

International Edition: DOI: 10.1002/anie.201612143 German Edition: DOI: 10.1002/ange.201612143

DNA-Encoded Compound Libraries as Open Source: **A Powerful Pathway to New Drugs**

Richard A. Lerner* and Sydney Brenner



Richard A. Lerner Sydney A. Brenner Institute Professor Professor The Scripps Research The Scripps Research Institute

Institute

For Academics https://delopen.org

DELopen Scientific Advisory Board

The board will set the direction and guide the development of DELopen in its vision to advance the adoption of DNA encoded library technology in new drug discovery.



Richard Lerner

Institute Professor Professor, Department of Chemistry, California



Phil Baran

Darlene Shiley Chair in Chemistry, Professor Department of Chemistry,



Carolyn Bertozzi

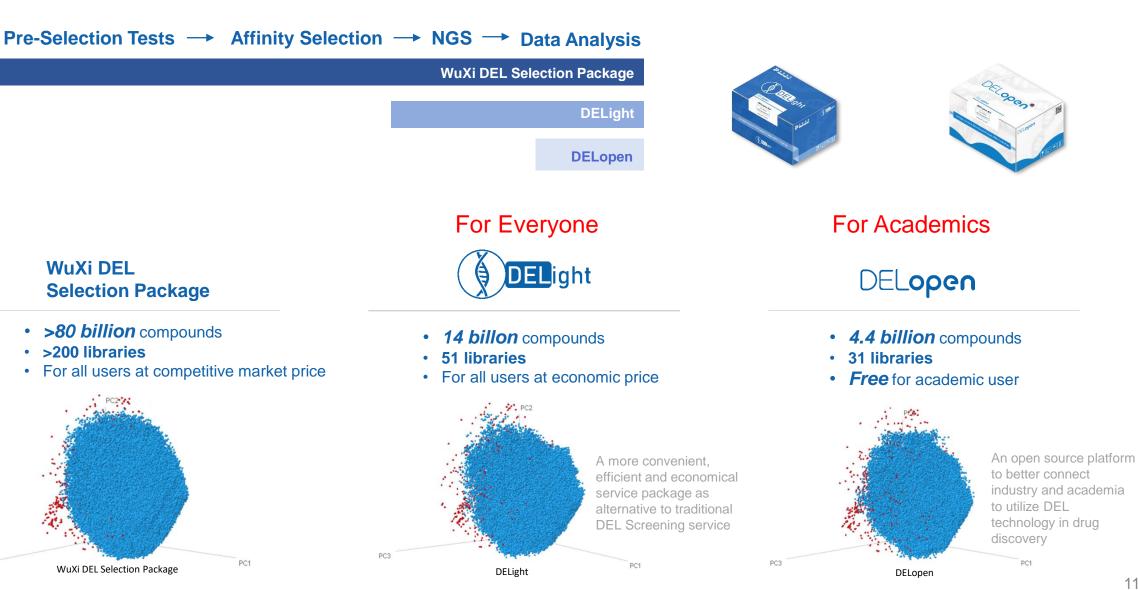
Director, ChEM-H, Anne T. and Robert M. Bass Professor in the School of



Raymond Dwek

Director of the Oxford Glycobiology Institute University of Oxford

WuXi DEL Screening Business Model



Proprietary building blocks (BBs)

PC3

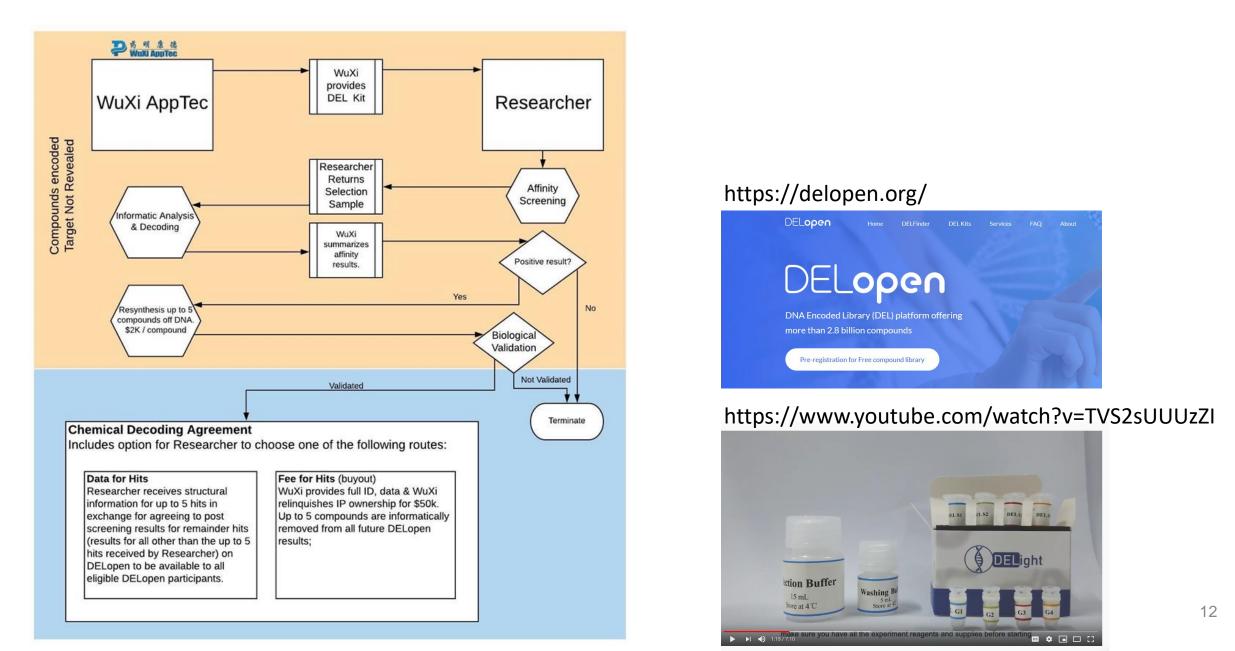
All commercial BBs

All commercial BBs

11

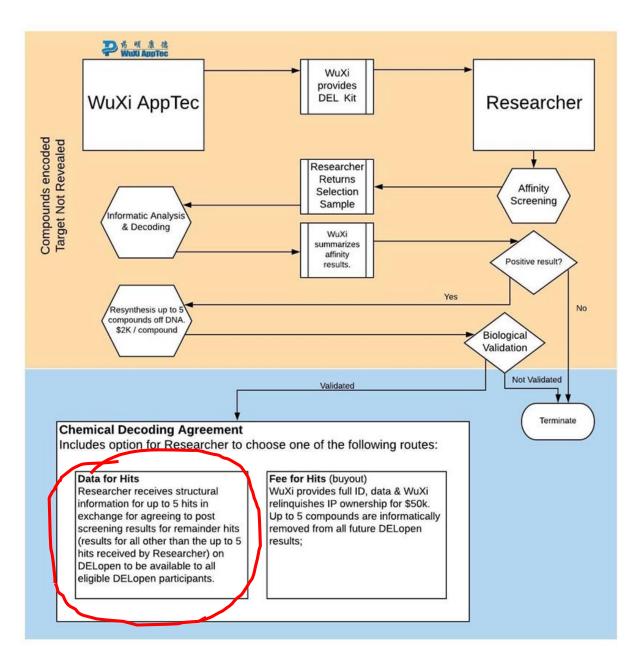
DELopen Workflow

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DELopen Workflow





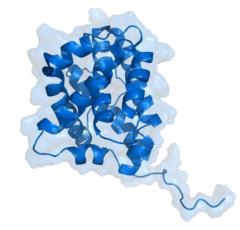
- You are free to work on your hits
 - patent
 - start a company
 - sell

"You Get What You Screen For"

- Frances Arnold, Nobel Laureate in Chemistry

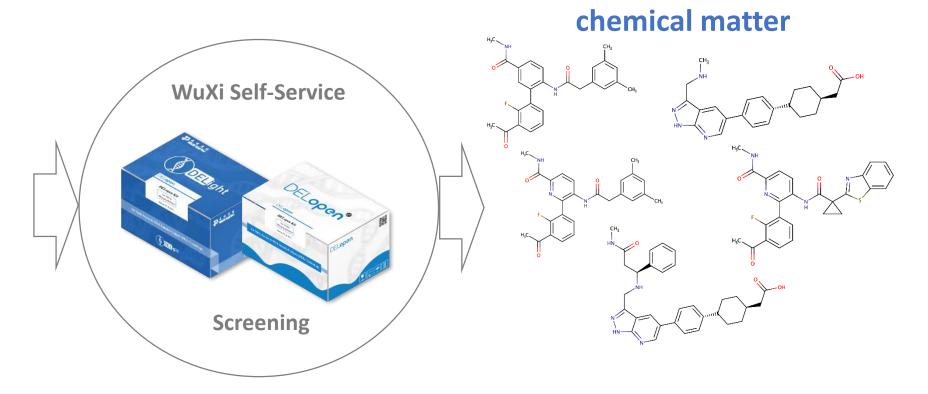


Potent drug-like



Target Quality

- >90% purity
- Not aggregated
- Correctly folded
 - Binding to positive control
 - Forming expected protein complexes





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