Antibiotic Discovery via BICyCLE (Biologically Inspired Chemically Created Leads)

A rapid, high-throughput drug discovery platform (BICyCLE) uses genetic information to quickly identify and validate promising natural product candidates, such as cyclic peptides, for developing new antibiotics against multi-drug resistant bacteria.

Researchers at Purdue University have developed a new drug discovery platform, Biologically Inspired Chemically Created Leads (BICyCLE) and have proven it through the discovery and validation of antibacterial cyclic peptides. New antibiotics are needed because bacteria responsible for serious infections continue to develop resistance to current treatments. Sixty-four percent of FDA-approved small molecule drugs are based on natural products, yet natural product discovery remains challenging. The biology of source organisms is often not amenable to scientific inquiry. A class of natural products of interest in pharmaceutical development are nonribosomal peptides (NRPs), cyclic molecules that are often bulky and structurally rigid, allowing them to favorably target permeable cell walls and resist proteases. Purdue researchers used genetic information to predict previously undiscovered NRPs. Through prediction and bioinformatic analysis, the researchers identified 131 unique cyclic peptides. Of those, 52 diverse peptides were amenable to synthesis. Fourteen of the synthesized compounds exhibited promising antibacterial activity in vitro, some with activity against multidrug-resistant Gram-negative bacteria.

Technology Validation: The new BICyCLE technique was used to establish 14 promising candidates for use in developing new antibiotics to treat multidrug resistant bacteria, synthesizing the NPs in over 90% purity.

Advantages:

- -High Throughput
- -Rapid Discovery of Bioactive Natural Products for Drug Discovery

Potential Applications:

Technology ID

2021-PARK-69383

Category

Biotechnology & Life
Sciences/Biomarker Discovery &
Diagnostics
Biotechnology & Life
Sciences/Bioinformatics &
Computational Biology
Pharmaceuticals/Drug Discovery
& Development
Biotechnology & Life
Sciences/Analytical & Diagnostic
Instrumentation
Pharmaceuticals/Small Molecule
Therapeutics
Pharmaceuticals/Research Tools
& Assays

Authors

Matthew Hostetler Elizabeth Ivy Parkinson

Further information

Joe Kasper JRKasper@prf.org

Nathan Smith nesmith@prf.org

View online



-Antibiotics Development

-Pharmaceutical Design

TRL: 3

Intellectual Property:

Provisional-Gov. Funding, 2021-03-04, United States | Utility-Gov. Funding, 2022-02-24, United States | CON-Gov. Funding, 2023-11-30, United States

Keywords: Drug discovery platform, BICyCLE, antibacterial cyclic peptides, new antibiotics, multidrug resistance, nonribosomal peptides, NRPs, bioinformatic analysis, high throughput drug discovery, pharmaceutical design