SERVICE GUIDE

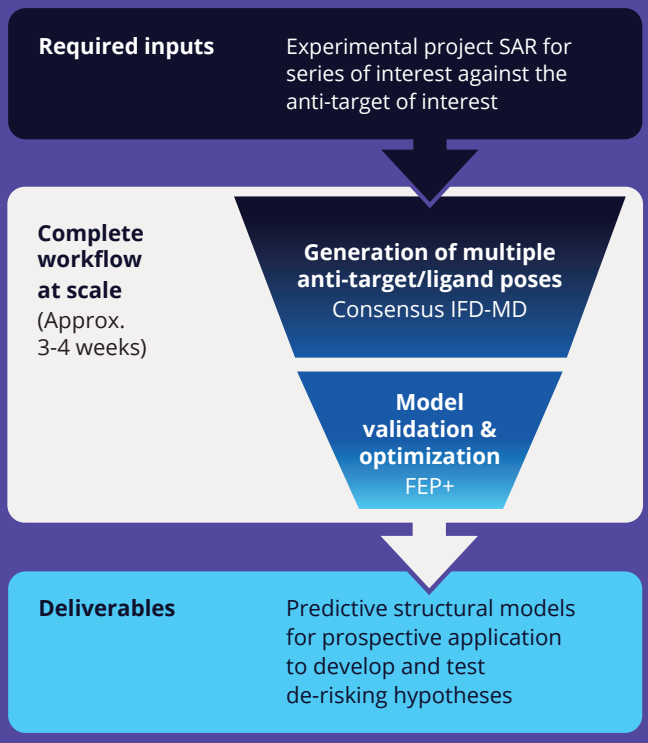
ADMET Liabilities Research Enablement Service

De-risk ADMET liabilities more efficiently using structure-based design

Resolve **CYP3A4**, **CYP2D6**, **hERG**, and **PXR** hurdles early to advance your drug discovery program. We'll help de-risk off-target liabilities by enabling FEP+ for common ADMET anti-targets, using a rigorous, structure-based approach powered by Schrödinger's technology and expertise.

Best suited for:

- Projects in early or advanced lead optimization that face a CYP, hERG, or PXR liability, including projects where prior efforts to resolve the liability have failed
- Companies who want to solve their ADMET issue by leveraging Schrödinger's industry-leading predictive modeling and large-scale computational power



Schrödinger

Advance your discovery program with unrivaled technologies and expertise

Case study: Prospective hERG modeling

Challenge
A project team within Schrödinger's therapeutics group discovered that lead compounds showed significant hERG inhibition. The team sought a structure-based approach to allow for rational, precision de-risking of hERG inhibition without impacting other project goals.

Result
A dataset of nine project compounds, each with an experimentally measured hERG IC50, were supplied. Using retrospective agreement for these nine compounds, a model that reproduced the physics of the system was identified and used to successfully rationalize prospective designs and eliminate the hERG liability.

Reference
Enabling structure-based drug discovery utilizing predicted models (Commentary). Miller EB, et al. *Cell*, 2024, 187, 3, 521-525.

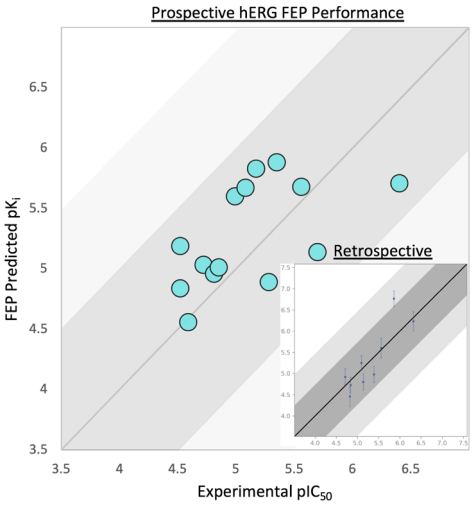


Figure: Prospective and retrospective FEP+ performance of hERG model.

Enabling digital technologies to drive discovery programs

FEP+

Digital assay for predicting protein-ligand binding across broad chemical space at an accuracy matching experimental methods

IFD-MD

Accurate ligand binding mode prediction for novel chemical matter, for on-targets and off-targets

Speak with an expert to get started

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