

# **'4318 (PKC-theta)** (in-licensed by BMY) Positive early Phase 1 results

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# **PKC-theta:** Positive early results from Phase 1 study

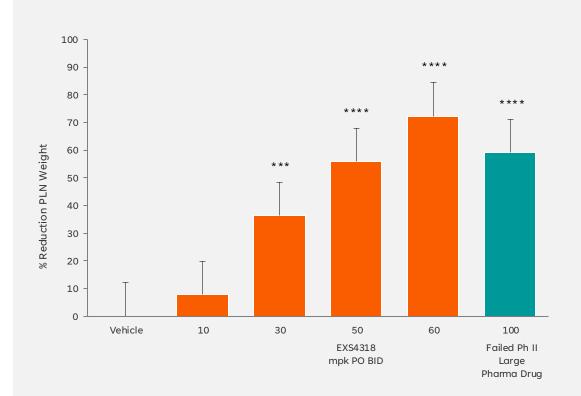
Expert led AI designed compound in-licensed by BMY, in August 2021

### **1** VVV Potential First-in-class Immunology Asset

- High-value immunology target that had **eluded many large biopharmas** due to selectivity challenges
- **Balanced profile** provided improvements in human whole blood potency and predicted human dose <200 mg/day
- Excellent selectivity versus near neighbours and broad kinome
- Positive early results from Ph1 study announced in May 2024

### Key Elements of TPP

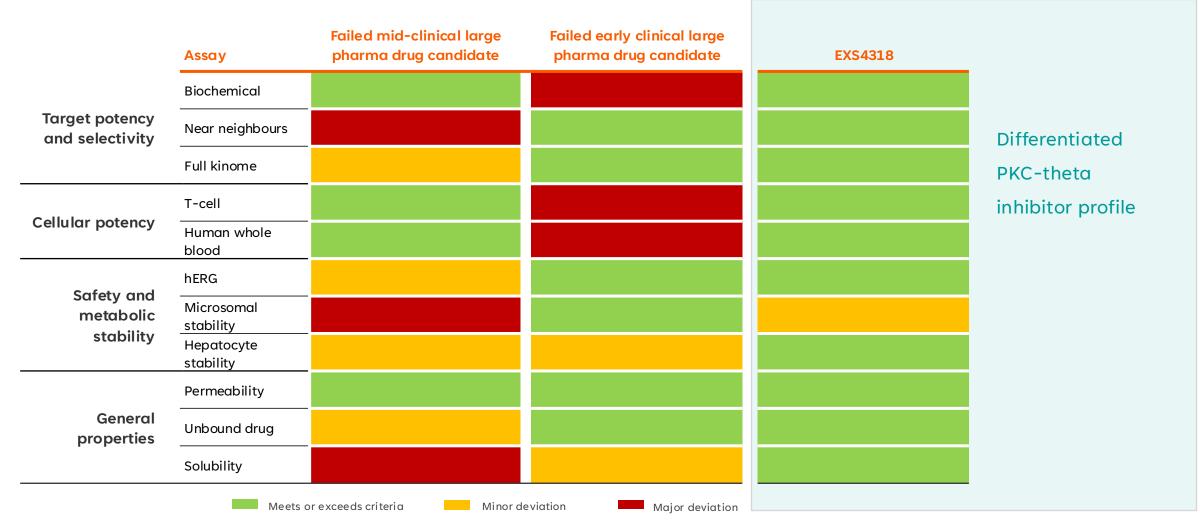
- 24 hrs coverage of IC<sub>80</sub> required to drive efficacy
- Predicted human dose <200 mg/day</li>
- High demands on target potency, selectivity, pharmacokinetics
- Robust translation into cellular and human whole blood assays



#### Better Efficacy at Approximately Half the Dose

# Large pharma failures on an attractive target

## Potential first-in-class immunology target



# Our approach

Fragments. 2D and 3D generative design. Hotspots and multi-task models



### Experiment

- Diverse ligand data sources.
  Proprietary fragment and kinase focussed SPR screens provided additional seed data
- Established and routinely executed key human whole blood assay

## **Expert-led AI Solutions**

- Generative design rapidly explored selectivity-focussed scaffolds; MERIT analysis quantified the most promising
- Hotspot and multi-task models drove local and global kinase selectivity, respectively

### **Best-in-class Compound**

- Nominated candidate designed in <11 months and was 150<sup>th</sup> novel compound prepared
- Demonstrates close relationship at Exscientia of AI and experiment
- Elegant solution to a challenging problem; Nominated candidate
   <400 MW</li>



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