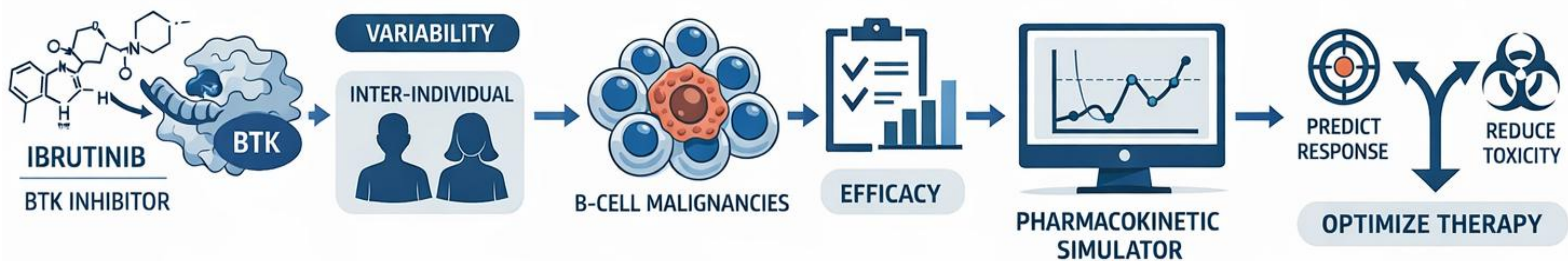


Development of a pharmacokinetic simulation tool for Ibrutinib to enable treatment individualization

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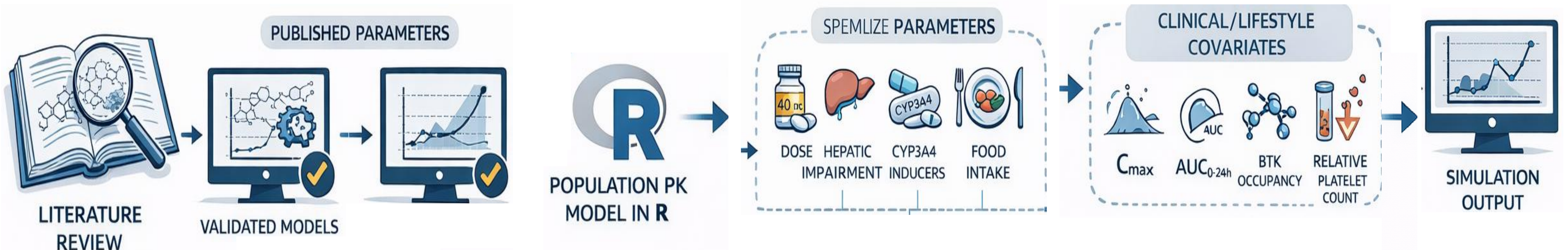
Background and Importance



Aim and Objectives

To develop a pharmacokinetic simulation tool for ibrutinib integrating cumulative modifiers of drug exposure, aiming to optimize therapeutic outcomes and support individualized treatment decisions.

Material and Methods



Results

Standard Conditions	
Parametro	Valor
C _{max}	23.6 ng/mL
AUC _{0-24h}	265.2 ng-h/mL
BTK Occupancy	>95%
Platelets	89.7% of baseline

Platelet reduction used as an indirect marker of hematological toxicity

In all scenarios, BTK occupancy >95%

Relative Change vs. Standard		
Factor/Condition	C _{max} Change	Platelet Reduction
Strong CYP3A4 Inhibitor	+172%	-18%
Moderate CYP3A4 Inhibitor	+65%	-7%
Moderate Hepatic Impairment	+45%	-5%
Food Intake	+49%	-5.7%
Smoking	-30%	NA*



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SIMULATIONS
VIRTUAL PATIENT CLINICAL SCENARIOS

Conclusions and Relevance

This population-based simulation tool accurately reproduced published pharmacokinetics and quantified the impact of clinical modifiers on ibrutinib exposure, efficacy, and safety. It provides a practical support for individualized dosing, improving therapeutic precision and reducing risk.

