Structural model

The logarithmic transformed concentration data were best described by a one-compartment model with an additive residual error. Inter-individual variability (IIV) was included on CL.

$CL_{i,j} = CL_{pop} * exp(\eta_i)$

Covariate analysis

Continuous covariates were tested using the following equation:

Eq. 1: $CL_{i,} = [CL_{pop} * (COV/median\ COV)^{\Theta cov}] * exp(\eta_i)$

Categorical covariates were implemented as shown in equation 2:

Eq. 2: $CL_{i,} = CL_{pop} * (\Theta_{cat1})^{FLAG1} * (\Theta_{cat2})^{FLAG2} * exp(\eta_i)$

The $\Theta_{cat1,2...}$ is the typical value under different categorical conditions The FLAG (1,2…) are first set to 0, when certain categorical covariate appears, the FLAG is set to 1.