

Table S2: Model parameters

| Parameter (unit) | Definition | Cyno Value | Human Value | Source |
|--|---|------------|-------------|---|
| MW _{FC} (Da) | Molecular weight of FLT3L-Fc | 83000 | 83000 | In-house measurement |
| MW _{CDX} (Da) | Molecular weight of CDX-301 | 35000 | 35000 | Ref. (1) |
| BW (kg) | Body weight | 2.6 | 70 | Nominal body weight value for species |
| V _{lymph} (L) | Volume of lymph | 0.086 | 5.2 | Cyno fit from data, human from (2) |
| V _{plasma} (L) | Volume of plasma | 0.09 | 2.6 | Cyno fit from data, human from (2) |
| CL _p (L/h) | Nonspecific clearance of FLT3L-Fc | 4.2E-04 | 6.94E-03 | Cyno fit from data, human allometrically scaled |
| CL _p (mL/d/kg) | Nonspecific clearance of FLT3L-Fc | 3.87 | 2.4 | Cyno fit from data, human allometrically scaled |
| σ_{leaky} | Leaky reflection coefficient | 0.656 | 0.656 | Fit from data |
| σ_{tight} | Tight reflection coefficient | 0.985 | 0.985 | Fit from data |
| σ_{lymph} | Lymph reflection coefficient | 0.2 | 0.2 | Ref. (2) |
| L (L/h) | Lymph flow rate | 0.012 | 0.121 | Cynomolgus monkey fit from data, human from (2) |
| K _p | Partition coefficient | 0.8 | 0.8 | Ref. (2) |
| ISF (mL/kg) | Interstitial fluid | 0.579 | 15.6 | Cynomolgus monkey fit from data, human from (3) |
| f_{leaky} | Fraction of leaky flow from blood to leaky | 0.65 | 0.65 | Ref. (2) |
| f_{tight} | Fraction of lymph flow from blood to tight | 0.35 | 0.35 | Ref. (2) |
| C _{R,0} (nM) | Initial concentration of FLT3 receptors in circulation | 11.3 | 4 | Fit from data |
| vm_{prolif} (h ⁻¹) | Maximum rate of FLT3 expansion induced by double-bound FLT3 receptor-drug complex | 1.90E-03 | 3.30E-3 | Fit from data |
| km_{prolif} | Amount of double-bound FLT3 receptor-drug complex to achieve 50% vm_{prolif} | 0.547 | 0.02 | Fit from data |
| α | Hill coefficient of FLT3 expansion induced by double-bound FLT3 receptor-drug complex | 3.93 | 1 | Fit from data |
| k _{deg} (h ⁻¹) | Turnover rate of FLT3 receptors in circulation | 0.016 | 0.016 | Fit from data |
| kon1 (1/nM/h) | On-rate for monovalent binding of drug to FLT3 receptors | 0.1 | 0.1 | Assumption |
| KD1 (nM) | monovalent binding affinity of drug to FLT3 receptors | 9 | 9 | Ref. (4) |
| kon2 (1/nM/h) | On-rate for binding of single-bound drug-FLT3 complex to the second FLT3 receptor | 3.6 | 3.6 | in-house Biacore measurement |
| KD2 (nM) | Binding affinity of single-bound drug-FLT3 complex to the second FLT3 receptor | 0.2 | 0.2 | in-house Biacore measurement |
| k _{absCDX} (h ⁻¹) | Absorption rate of CDX-301 from SC to plasma | NA | 0.04 | Fit from CDX-301 data (5) |

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|--|---|----|----------|------------------------------|
| $V_{d_{CDX}}$ (mL/kg) | Volume of distribution of CDX-301 | NA | 213 | Fit from CDX-301 data (5) |
| $v_{m_{CDX}}$ (mL/h/kg) | Maximum nonlinear elimination rate of CDX-301 | NA | 3.72 | Fit from CDX-301 data (5) |
| $k_{m_{CDX}}$ ($\mu\text{g/mL}$) | Concentration of CDX-301 needed to achieve 50% of $V_{m_{CDX}}$ | NA | 0.394 | Fit from CDX-301 data (5) |
| $init_{DC1}$ (cells/mL) | Mean initial cDC1 count | NA | 1180 | Human estimated from (5) |
| $k_{deg_{DC1}}$ (h^{-1}) | Turnover rate of the cDC1 population | NA | 1.60E-02 | Fit from CDX-301 data (5, 6) |
| $v_{m_{DC1}}$ (h^{-1}) | Maximum rate of cDC1 expansion induced by free drug | NA | 1.61 | Fit from CDX-301 data (5, 6) |
| $k_{m_{DC1}}$ (nM) | Concentration of free drug to increase cDC1 counts to 50% $V_{m_{DC1}}$ | NA | 0.072 | Fit from CDX-301 data (5, 6) |
| del_{DC1} (h^{-1}) | Transit rate between cDC1 compartments | NA | 0.114 | Fit from CDX-301 data (5, 6) |
| $n_{1_{DC1}}$ | Hill coefficient of free drug effect on cDC1 expansion | NA | 5.85 | Fit from CDX-301 data (5, 6) |
| $init_{DC2}$ (cells/mL) | Mean initial cDC2 count | NA | 12700 | Human estimated from (6) |
| $k_{deg_{DC2}}$ (h^{-1}) | Turnover rate of the cDC2 population | NA | 1.70E-03 | Fit from CDX-301 data (5, 6) |
| $v_{m_{DC2}}$ (h^{-1}) | Maximum rate of cDC2 expansion induced by free drug | NA | 16.1 | Fit from CDX-301 data (5, 6) |
| $k_{m_{DC2}}$ (nM) | Concentration of free drug to increase cDC2 counts to 50% $V_{m_{DC2}}$ | NA | 0.209 | Fit from CDX-301 data (5, 6) |
| del_{DC2} (h^{-1}) | Transit rate between cDC2 compartments | NA | 0.053 | Fit from CDX-301 data (5, 6) |
| $n_{1_{DC2}}$ | Hill coefficient of free drug effect on cDC2 expansion | NA | 0.888 | Fit from CDX-301 data (5, 6) |
| $k_{deg_{2_{DC}}}$ (h^{-1}) | Second-order apoptosis rate of cDC2 | NA | 3.64E-04 | Fit from CDX-301 data (5, 6) |
| f_{DC1} | Fractional rate of cDC1 second-order apoptosis rate compared to that of DC2 | NA | 0.476 | Fit from CDX-301 data (5, 6) |

References

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