

Table S1: Equations for kinetic model of ^{64}Cu signal in the kidney arising from liposomal ^{64}Cu

| State Variable | Equation |
|---|--|
| Total volume, kidney | $V_{kidney,total} = V_{cap} + V_{tissue}$ |
| ^{64}Cu concentration, blood | $\frac{d[Lipo64\text{Cu}]_{blood}}{dt} = -k_{el} \cdot [Lipo64\text{Cu}]_{blood} - Q \cdot [Lipo64\text{Cu}]_{blood} + Q \cdot [Lipo64\text{Cu}]_{cap} \cdot \frac{V_{cap}}{V_{blood}}$ |
| ^{64}Cu concentration, kidney vascular space | $\frac{d[Lipo64\text{Cu}]_{cap}}{dt} = Q \cdot [Lipo64\text{Cu}]_{blood} \cdot \frac{V_{blood}}{V_{cap}} - Q \cdot [Lipo64\text{Cu}]_{cap} - k_{in} \cdot [Lipo64\text{Cu}]_{cap} + k_{out} \cdot [Lipo64\text{Cu}]_{tissue} \cdot \frac{V_{tissue}}{V_{cap}}$ |
| ^{64}Cu concentration, kidney tissue | $\frac{d[Lipo64\text{Cu}]_{tissue}}{dt} = k_{in} \cdot [Lipo64\text{Cu}]_{cap} \cdot \frac{V_{cap}}{V_{tissue}} - k_{out} \cdot [Lipo64\text{Cu}]_{tissue} - k_{rpu} \cdot [Lipo64\text{Cu}]_{tissue}$ |
| ^{64}Cu concentration, renal pelvis | $\frac{d[64\text{Cu}]_{rp}}{dt} = k_{rpu} \cdot [Lipo64\text{Cu}]_{tissue} \cdot \frac{V_{tissue}}{VF_{rp} \cdot V_{kidney,total}}$ |
| ^{64}Cu total signal, total kidney | $64\text{Cu}_{kidney,total} = [Lipo64\text{Cu}]_{cap} \cdot V_{cap} + [Lipo64\text{Cu}]_{tissue} \cdot V_{tissue} + [64\text{Cu}]_{rp} \cdot VF_{rp} \cdot V_{kidney,total}$ |
| ^{64}Cu total signal, renal pelvis | $64\text{Cu}_{rp,total} = [64\text{Cu}]_{rp} \cdot VF_{rp} \cdot V_{kidney,total} + ([Lipo64\text{Cu}]_{cap} \cdot V_{cap} + [Lipo64\text{Cu}]_{tissue} \cdot V_{tissue})VF_{rp}$ |

The model was implemented in MATLAB SimBiology R2014A (The Mathworks, Natick, MA). For questions, please contact Bart Hendriks (bhendriks@merrimackpharma.com).

Table S2: Kinetic Model Parameter Descriptions

| Parameter | Description | Monkey Value | Human Value | Units | Reference |
|--------------------|--|------------------------------|------------------------------|-------|---|
| V_{blood} | Blood volume | 0.1 | 4.875 | L | Estimated from 1-compartment PK estimates using imaging data; Lee, Lan Na (1998). "Volume of Blood in a Human". The Physics Factbook. |
| $V_{kidney,total}$ | Total kidney volume | 0.00487 | 0.31 | L | Measured from imaging data; PMID: 8378254* |
| V_{cap} | Kidney capillary volume | $0.34 * V_{kidney,total}$ | $0.34 * V_{kidney,total}$ | L | NA |
| V_{tissue} | Kidney tissue volume | $V_{kidney,total} - V_{cap}$ | $V_{kidney,total} - V_{cap}$ | L | NA |
| VF_{rp} | Fraction of total kidney volume that comprises the renal pelvis | 0.1 | 0.1 | - | Estimated from data |
| Q | Blood flow rate into/out of kidneys | $5.52 * V_{kidney,total}$ | $4 * V_{kidney,total}$ | L/min | PMID: 8378254 |
| k_{el} | Elimination rate constant for liposome clearance from blood | 0.000467 | 0.000218 | 1/min | Estimated from data; Doxil package insert |
| k_{in} | Rate constant for liposomal deposition from kidney capillaries into kidney tissue | 2.55E-03 | 2.55E-03 | 1/min | Estimated from data |
| k_{out} | Rate constant for liposomal washout from kidney tissue into kidney capillaries | 1.00E-03 | 1.00E-03 | 1/min | Estimated from data |
| k_{rp} | Rate constant for renal pelvis uptake of ^{64}Cu from tissue deposited liposomal ^{64}Cu | 2.64E-04 | 2.64E-04 | 1/min | Estimated from data |

*Tissue density was assumed to be 1 kg/L.

Squirrel monkey weight was assumed to be 0.5 kg.