



Supplemental Figure 2. Plasma protein binding. (A) Correlation between total and unbound quinidine concentrations in plasma from the brain distribution experiments. There is a strong linear correlation between total and unbound concentrations ($r^2=0.953$, $N=24$ samples, $p=0.000$). (B and C) Effect of tariquidar treatment (B, $n=12$ samples per group) and experimental time point (C, $n=4$ samples per group) on the unbound fraction of quinidine in plasma. A two-way ANOVA showed no significant effect of TQD treatment (TQD vs. vehicle: $F(1,17)=1.70$, $p=0.21$) and of time of administration ($F(5,17)=2.52$, $p=0.07$). (D) Effect of experimental set-up on the unbound fraction of quinidine in plasma. There is no significant difference in the unbound fraction between brain distribution (projA, $n=24$ samples) and microdialysis experiments (projB, $n=7$ samples) ($t(29)=0.747$, $p=0.46$, t-test). Mean \pm SEM in all graphs. N.s.: not significant.