#### Supplemental Data for Molecular Pharmacology article:

## New Positive K<sub>Ca</sub> Channel Gating Modulators with Selectivity for K<sub>Ca</sub>3.1

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#### This file contains:

- 1 Supplemental Figure showing the hydrogen bonding pattern of SKA-120 and SKA-121
- 1 Supplemental Figure showing QPatch recordings of Na<sub>V</sub>1.2 and Na<sub>V</sub>1.5 currents
- 1 Supplemental Figure showing telemetric blood pressure measurements after i.p. injection of 30 mg/kg SKA-111, SKA-121

Hydrogen bonding scheme for SKA-120

Hydrogen bonding scheme for SKA-121

# **Supplemental Figure 1:**

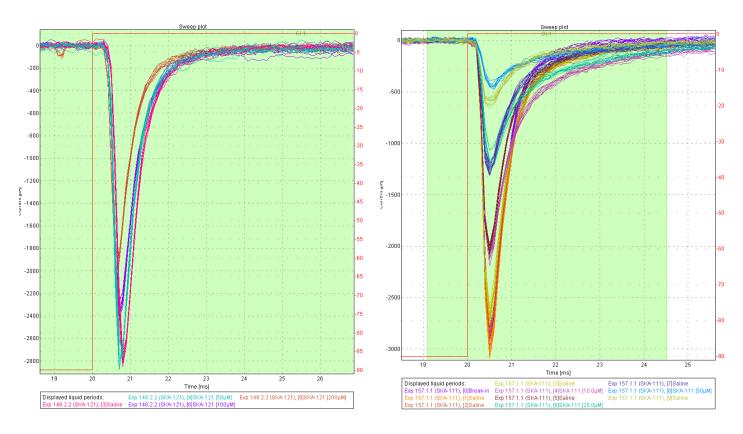
Differences in the hydrogen bonding pattern in the crystal structures of SKA-120 and SKA-121.

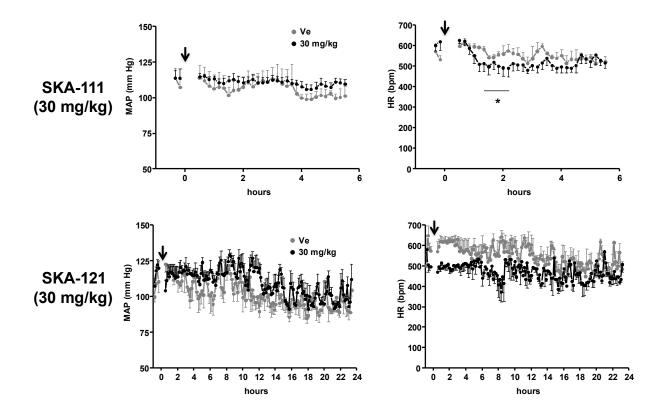
# **Supplemental Figure 2:**

Representative sweep plots of QPatch  $Na_V1.2$  recordings from N1E-115 neuroblastoma and  $Na_V1.5$  recordings from HEK-293 cells. Currents were elicited by 20 ms pulses from -90 mV to 0 mV applied every 10 sec with a KF-based internal solution and normal Ringer as an external solution.



Na<sub>V</sub>1.5 (HEK-293 cells)





### **Supplemental Figure 3:**

Telemetric blood pressure measurements after i.p. injection of 30 mg/kg SKA-111, SKA-121, or vehicle (Ve). Note that only SKA-111 moderately reduced heart rate (HR) at 2 h after injection while SKA-111 or SKA-121 did not change mean arterial pressure (MAP). Data points are means  $\pm$  SEM; n = 3-4 per group. Line in right panel on top indicates time period when HR was significantly different from Ve. \*<0.05, unpaired Student T-test.