

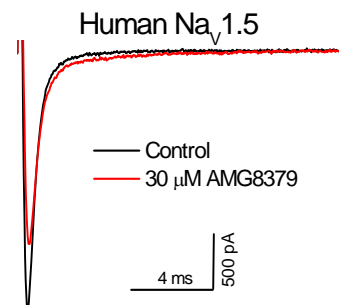
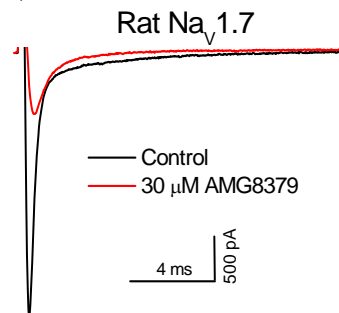
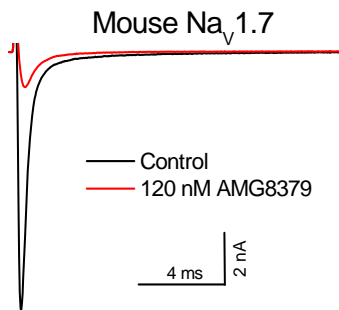
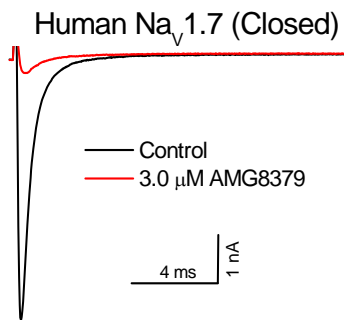
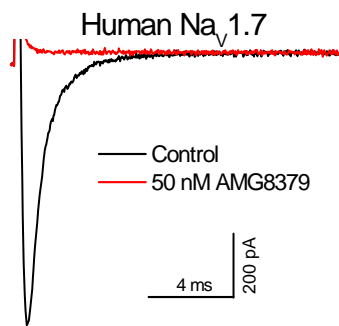
Supplementary Information

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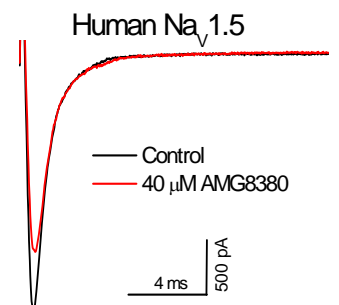
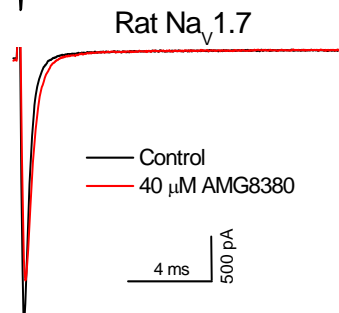
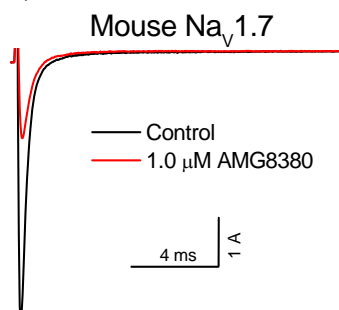
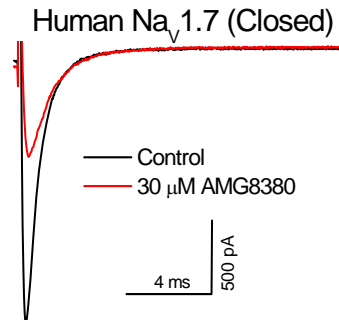
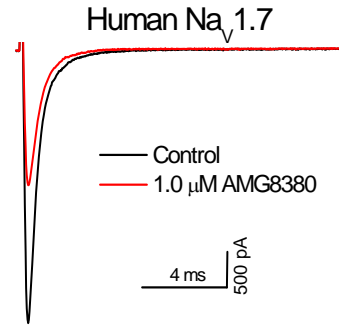
Pharmacologic Characterization of AMG8379, a Potent and Selective Small Molecule Sulfonamide Antagonist of the Voltage-Gated Sodium Channel Na_v1.7

Thomas Kornecook, Ruoyuan Yin, Stephen Altmann, Xuhai Be, Virginia Berry, Christopher P. Ilch, Michael Jarosh, Danielle Johnson, Josie H. Lee, Sonya G. Lehto, Joseph Ligutti, Dong Liu, Jason Luther, David Matson, Danny Ortuno, John Roberts, Kristin Taborn, Jinti Wang, Matthew M. Weiss, Violeta Yu, Dawn X. D. Zhu, Robert T. Fremeau Jr., and Bryan D. Moyer

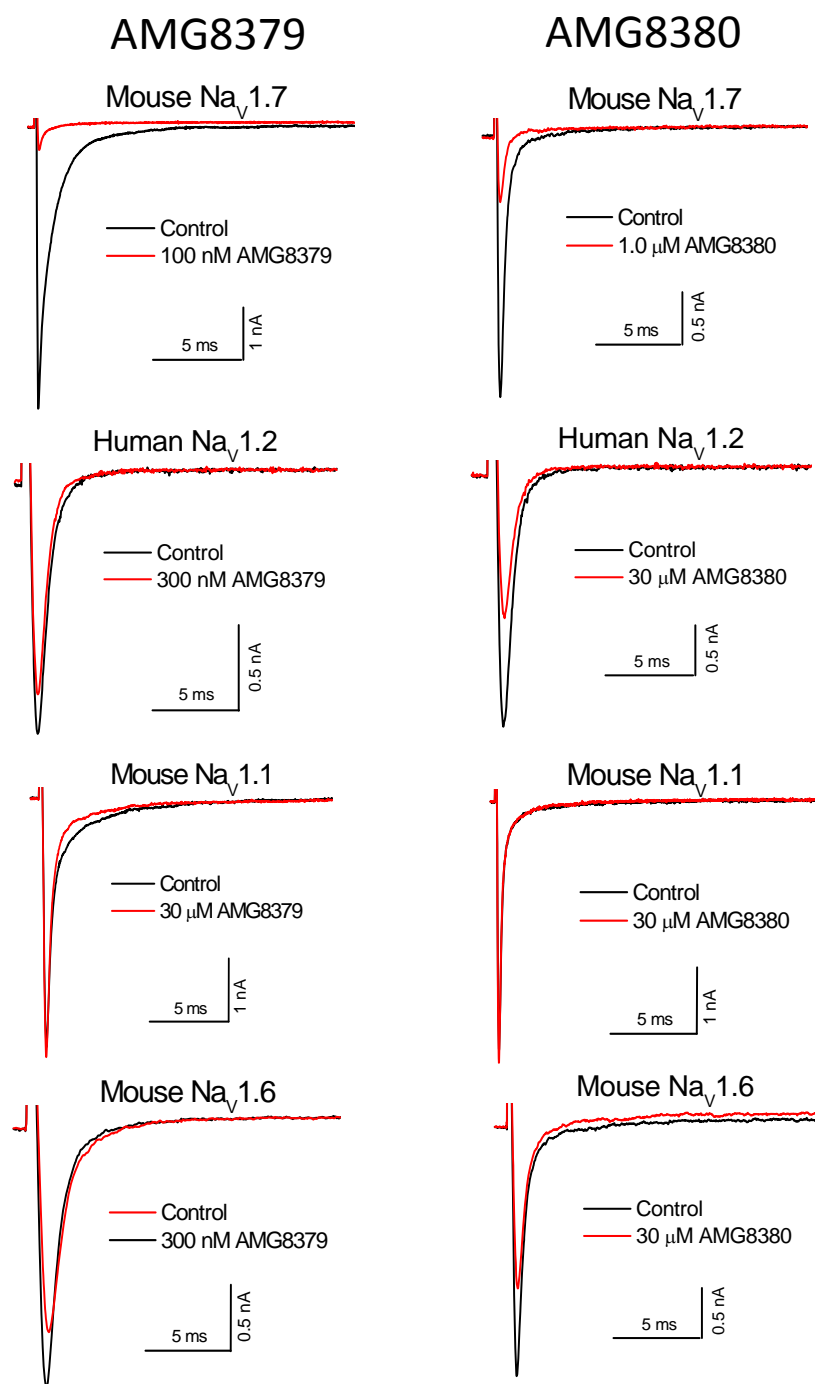
AMG8379



AMG8380



Supplementary Figure 1. Representative current traces illustrating potent and selective block of hNa_v1.7 and mNa_v1.7 but not rNa_v1.7 or hNa_v1.5 in a partially inactivated state or hNa_v1.7 channels in the resting/closed state by AMG8379 compared to AMG8380 in heterologous cells. Patch clamp recordings were from the PatchXpress platform. Traces show currents for the indicated channel before (black) and after (red) compound addition at the indicated concentrations.



Supplementary Figure 2. Representative current traces illustrating potent and selective block of mNa_v1.7, but not hNa_v1.2, mNa_v1.1, and mNa_v1.6, channels by AMG8379 compared to AMG8380 in heterologous cells. Manual patch clamp electrophysiology recordings used the IonWorks Quattro voltage protocol. Traces show currents for the indicated channel before (black) and after (red) compound addition at the indicated concentrations.