

Supplementary Table S1. Currents and $V_{1/2}$ in homomeric and heteromeric transfected HEK293 cell in the variants of p.Gly281Glu, p.Thr287Ile, and p.Gly281Glu.

	N	Currents at -10 mV (pA) (mean±SD)	Currents at 0 mV (pA) (mean±SD)	Currents at +10 mV (pA) (mean±SD)	Currents at +20 mV (pA) (mean±SD)	Currents at +30 mV (pA) (mean±SD)	Currents at +40 mV (pA) (mean±SD)
<i>KCNQ2</i> WT (2 ug) *	22	596.5 ± 121.5	769.3 ± 119.3	954.1 ± 121.5	1145.8 ± 117.9	1333.9 ± 120.8	1534.5 ± 141.9
p.Gly281Glu (2 ug)	8	480.9 ± 101.5 t(30) = 3.51, p = 0.001	600.0 ± 96.6 t(30) = 4.70, p < .001	721.8 ± 89.3 t(30) = 6.07, p < .001	852.8 ± 91.6 t(30) = 7.25, p < .001	979.6 ± 97.1 t(30) = 8.78, p < .001	1109.4 ± 112.6 t(30) = 8.76, p < .001
p.Thr287Ile (2 ug)	10	505.3 ± 56.25 t(30) = 2.25, p = .032	649.9±85.7 t(30) = 2.84, p = .008	789.5±92.9 t(30) = 3.80, p = .001	937.7 ± 99.1 t(30) = 4.85, p < .001	1087.0 ± 107.3 t(30) = 5.54, p < .001	1240.1 ± 122.2 t(30) = 5.66, p < .001
p.Pro285Thr (2 ug)	10	465.2 ± 64.0 t(30) = 3.20, p = .003	599.7±76.4 t(30) = 4.11, p < .001	747.5±93.7 t(30) = 4.76, p < .001	890.9 ± 110.5 t(30) = 5.78, p < .001	1035 ± 110.6 t(30) = 6.65, p < .001	1173.3 ± 126.5 t(30) = 6.89, p < .001
<i>KCNQ2</i> WT + p.Gly281Glu (1ug:1 ug)	8	534.0 ± 75.3 t(30) = 1.43, p = .164	679.1±61.0 t(30) = 2.14, p = .041	843.4±63.6 t(30) = 2.57, p = .015	987.1 ± 51.0 t(30) = 3.86, p < .001	1137.6 ± 64.3 t(30) = 4.59, p < .001	1323.1 ± 81.0 t(30) = 4.18, p < .001
<i>KCNQ2</i> WT + p.Thr287Ile (1ug:1 ug)	10	548.3 ± 62.37 t(30) = 1.18, p = .248	696.0±76.3 t(30) = 1.78, p = .086	837.5±88.0 t(30) = 2.72, p = .011	1007.6 ± 95.3 t(30) = 3.25, p = .003	1164.3 ± 112.8 t(30) = 3.75, p < .001	1323.5 ± 99.9 t(30) = 4.23, p < .001
<i>KCNQ2</i> WT + p.Pro285Thr (1ug:1 ug)	10	540.9 ± 76.04 t(18) = 1.33, p = 0.194	687.2±90.3 t(18) = 1.93, p = 0.063	858.8±102.6 t(18) = 2.15, p = 0.040	1030.2 ± 122.6 t(18) = 2.54, p = 0.016	1206.8 ± 152.5 t(18) = 2.54, p = 0.016	1385.4 ± 163.6 t(30) = 2.63, p = 0.013
<i>KCNQ2</i> WT + <i>KCNQ3</i> WT (1ug:1 ug) §	10	934.6 ± 151.96	1218.5±184.3	1530.9±236.7	1843.1 ± 290.9	2168.3 ± 330.6	2474.3 ± 395.3
<i>KCNQ2</i> WT + p.Gly281Glu + <i>KCNQ3</i> WT (0.5 ug: 0.5 ug: 1 ug)	8	673.4 ± 87.4 t(18) = 4.31, p = 0.001	914.1±120.9 t(18) = 4.02, p = 0.001	1179.1±129.6 t(18) = 3.76, p = 0.002	1481.4±116.6 t(18) = 3.30, p = 0.005	1824.5 ± 117.5 t(18) = 2.79, p = 0.013	2120.0 ± 65.9 t(18) = 2.49, p = 0.024
<i>KCNQ2</i> WT + p.Thr287Ile + <i>KCNQ3</i> WT	10	818.1 ± 185.8	1012.8±211.6	1244.7±265.0	1435.4±218.4	1667.1 ± 225.3	1865.0 ± 155.9

(0.5 ug: 0.5 ug: 1 ug)		t(18) = 1.50, <i>p</i> = 0.150	t(18) = 2.38, <i>p</i> = 0.029	t(18) = 2.63, <i>p</i> = 0.017	t(18) = 3.43, <i>p</i> = 0.003	t(18) = 3.86, <i>p</i> = 0.001	t(18) = 4.24, <i>P</i> < .001
<i>KCNQ2</i> WT + p.Pro285Thr + <i>KCNQ3</i> WT	10	820.9 ± 133.1	1065.8 ± 144.9	1321.1 ± 165.1	1570.0 ± 190.6	1803.6 ± 223.6	2038.8 ± 251.4
(0.5 ug: 0.5 ug: 1 ug)		t(18) = 1.78, <i>p</i> = 0.092	t(18) = 2.06, <i>p</i> = 0.054	t(18) = 2.30, <i>p</i> = 0.034	t(18) = 2.48, <i>p</i> = 0.023	t(18) = 2.89, <i>p</i> = 0.010	t(18) = 2.94, <i>p</i> = 0.009

Bold fonts indicate *p* < 0.05.

WT, wild type; SD, standard deviation. The data were then fit to a Boltzmann distribution of the following form: $G/G_{max} = 1/(1 + \exp[(V - V_{1/2})/dx])$.

The currents in the homomeric transfected variants and heteromeric transfected *KCNQ2* WT + variants were compared with the currents in *KCNQ2* WT respectively.

\$ The currents in the heteromeric *KCNQ2* WT + *KCNQ3* WT + variants were compared with the currents in *KCNQ2* WT + *KCNQ3* WT respectively.