

Voltage-dependent activation in EAG channels follows a ligand-receptor rather than a mechanical-lever mechanism

**Olfat A. Malak^{1#}, Grigory S. Gluhov², Anastasia V. Grizel³, Kseniya S. Kudryashova^{2,4},
Olga S. Sokolova^{2,§}, Gildas Loussouarn^{1,§*}**

Running Title: Voltage-dependent gating mechanism of EAG channels

present address: Cardiovascular Institute, Stanford University, 300 Pasteur Drive, Stanford, CA 94305-5111

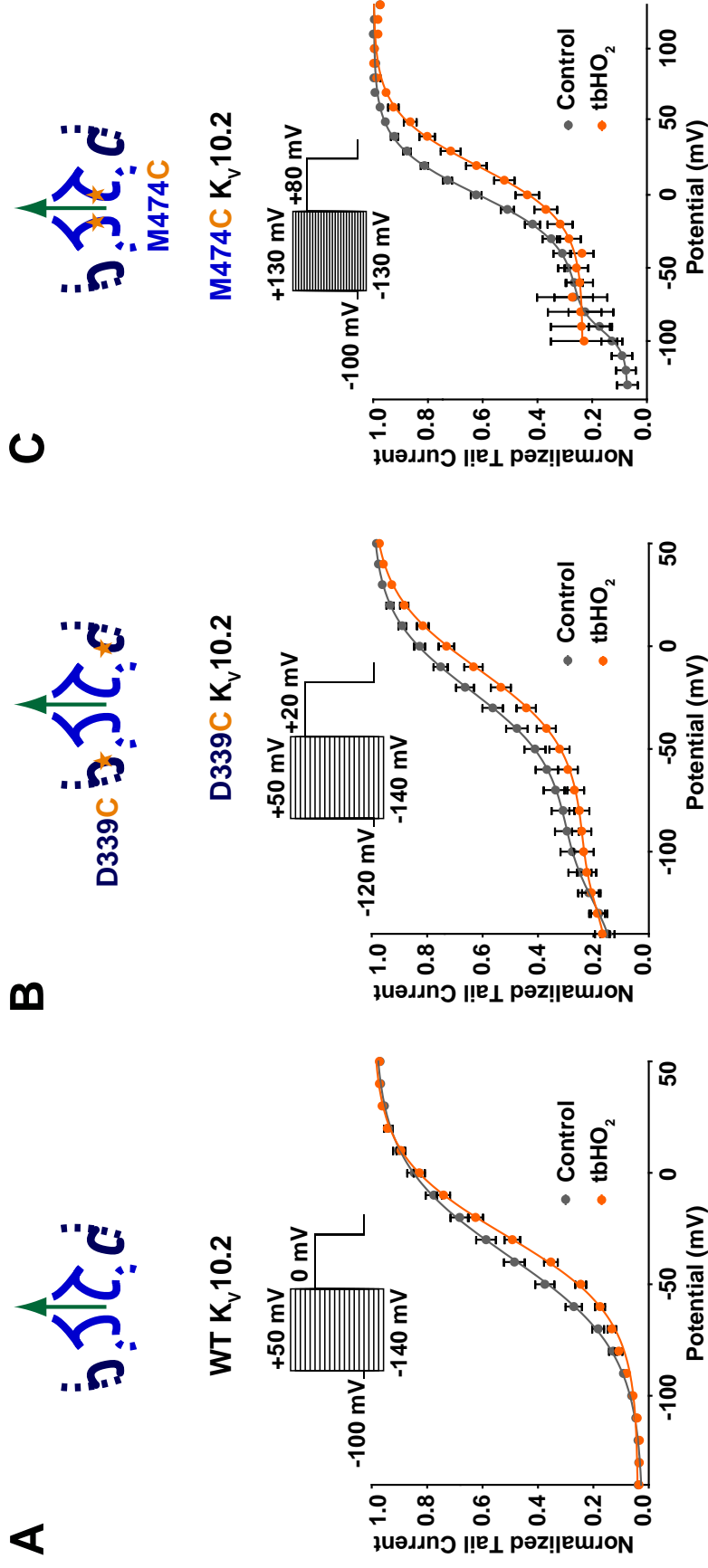
*From the ¹INSERM, CNRS, l'Institut du Thorax, Université de Nantes, 44007, Nantes, France ;
²Moscow M.V. Lomonosov State University, Moscow 119234, Russia; ³Saint Petersburg State University, Saint Petersburg 199034, Russia; ⁴Shemyakin-Ovchinnikov Institute of Bioorganic Chemistry of Russian Academy of Sciences, Moscow, 117997, Russia*

§ These authors contributed equally to this work.

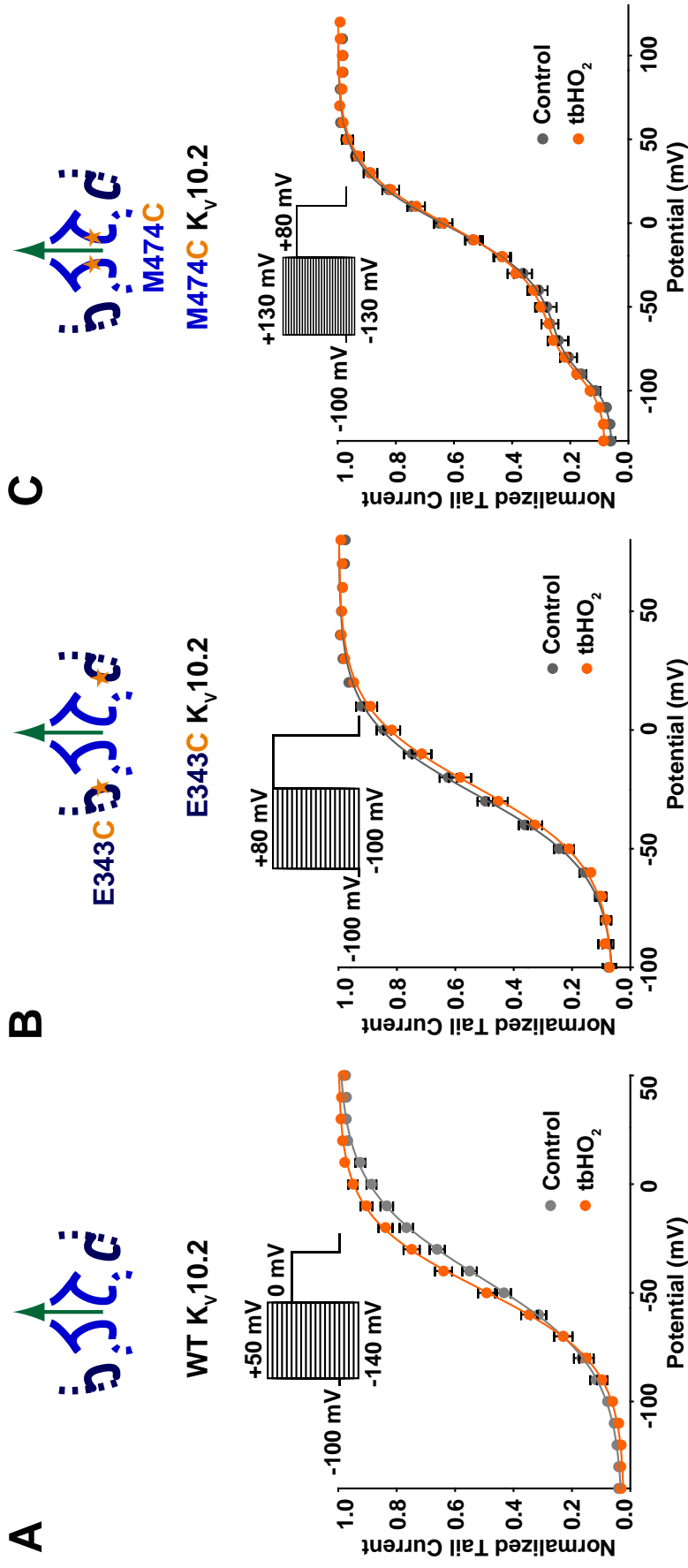
**To whom correspondence should be addressed: Dr. Gildas Loussouarn, L'institut du thorax, INSERM UMR 1087 / CNRS UMR 6291, IRS-UN, 8 Quai Moncouso BP 70721, 44007 Nantes cedex 1, France; gildas.loussouarn@inserm.fr; Tel +33 (0)2 2808 0150*

Keywords: EAG channel, Kv10.2 channel, voltage dependence, S4-S5 linker, S6 C-terminus, N-CAP, PAS domain, allostery, voltage-gated cation channel

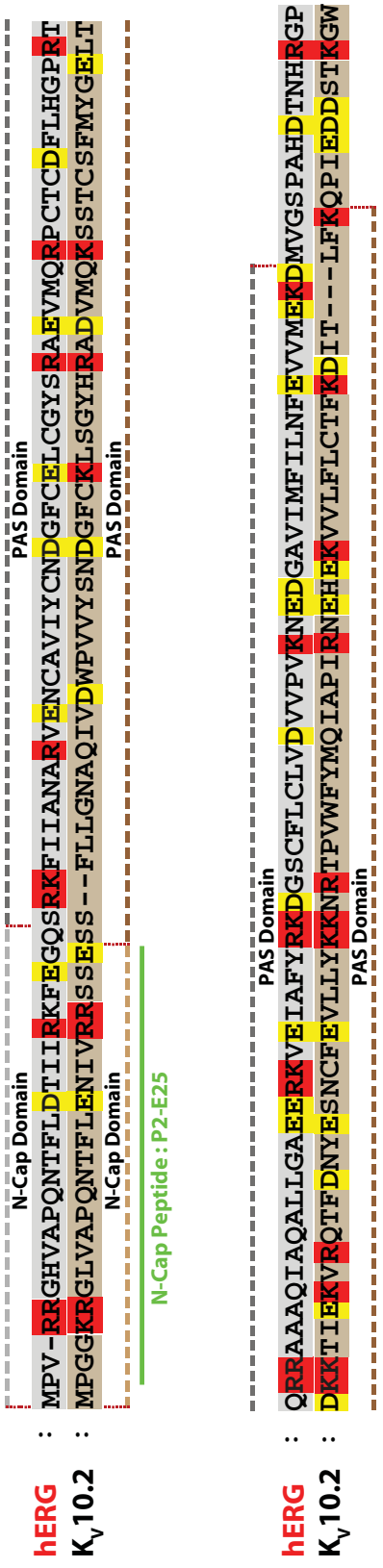
SUPPLEMENTAL FILE



Supplemental figure 1: activation curve in WT, D339C and M474C $K_v10.2$, in presence and absence of 0.2 mM tbHO₂ for 2h. A: Activation curve of WT $K_v10.2$, obtained from recordings as in Figure 2C, after 2h incubation in Tyrode without (control) or with 0.2 mM tbHO₂ (tbHO₂). **B:** Activation curve of D339C $K_v10.2$, obtained from recordings as in Figure 2E, after 2h incubation in Tyrode without (control) or with 0.2 mM tbHO₂ (tbHO₂). **C:** Activation curve of M474C $K_v10.2$, obtained from recordings as in Figure 2G, after 2h incubation in Tyrode without (control) or with 0.2 mM tbHO₂ (tbHO₂).



Supplemental figure 2: activation curve in WT, E343C and M474C $K_v10.2$, in presence and absence of 2 mM tbHO₂ for 15 min. A: Activation curve of WT $K_v10.2$, obtained from recordings as in Figure 4C, after 15 minutes incubation in Tyrode without (control) or with 2 mM tbHO₂ (tbHO₂). **B:** Activation curve of E343C $K_v10.2$, obtained from recordings as in Figure 4E, after 15 minutes incubation in Tyrode without (control) or with 2 mM tbHO₂ (tbHO₂). **C:** Activation curve of M474C $K_v10.2$, obtained from recordings as in Figure 4G, after 15 minutes incubation in Tyrode without (control) or with 2 mM tbHO₂ (tbHO₂).



Supplemental figure 3: K_v10.2 N-Cap and PAS domains. The alignment was obtained using Cobalt (Papadopoulos and Agarwala, 2007).