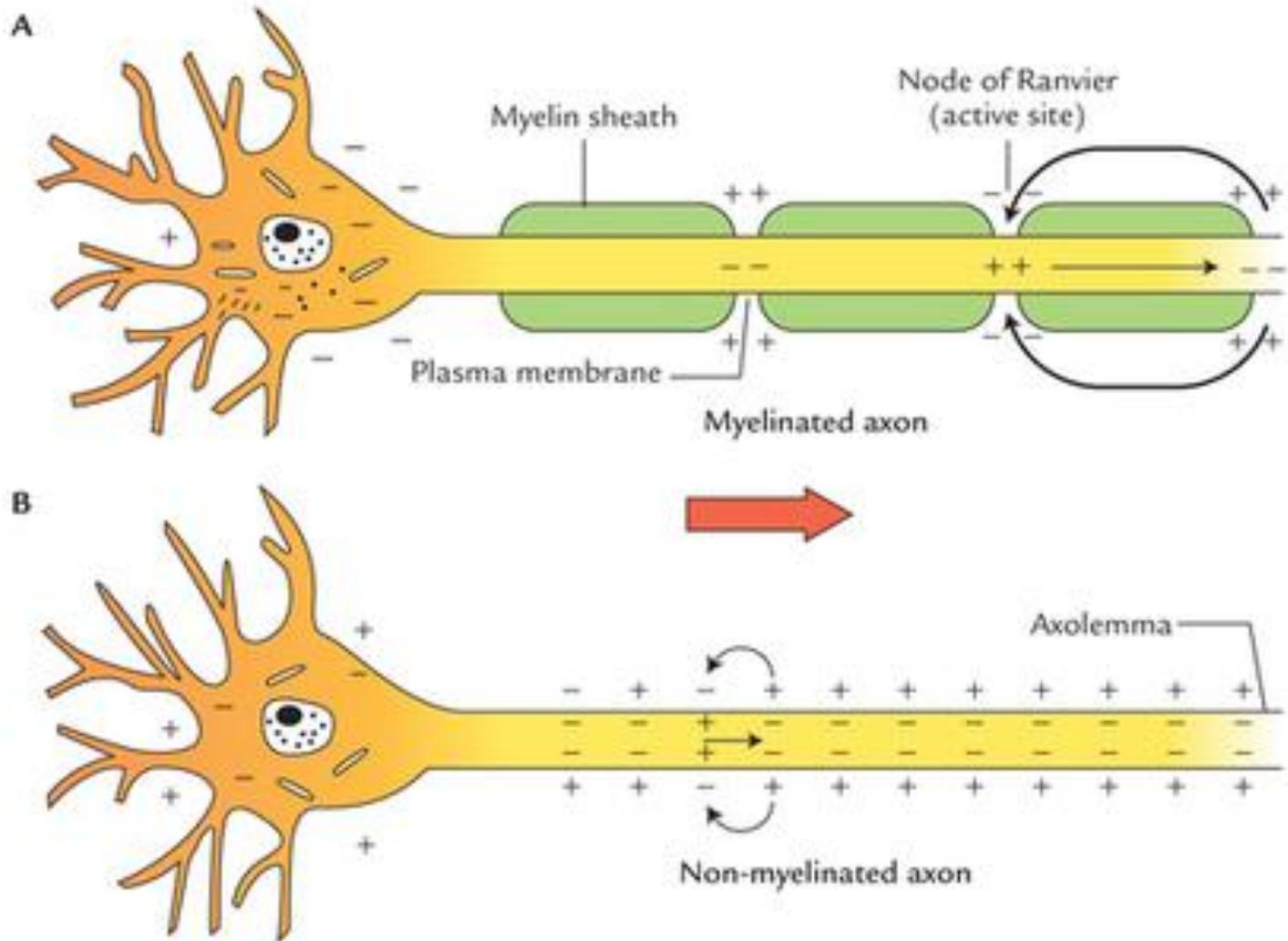


$K_V$ ,  $Na_V$ ,  $Ca_V$  are voltage gated ion channels

NCX is the  $Na^+-Ca^{2+}$  exchanger

$K_{(Ca)}$  is the  $Ca^{2+}$  activated  $K^+$  channel



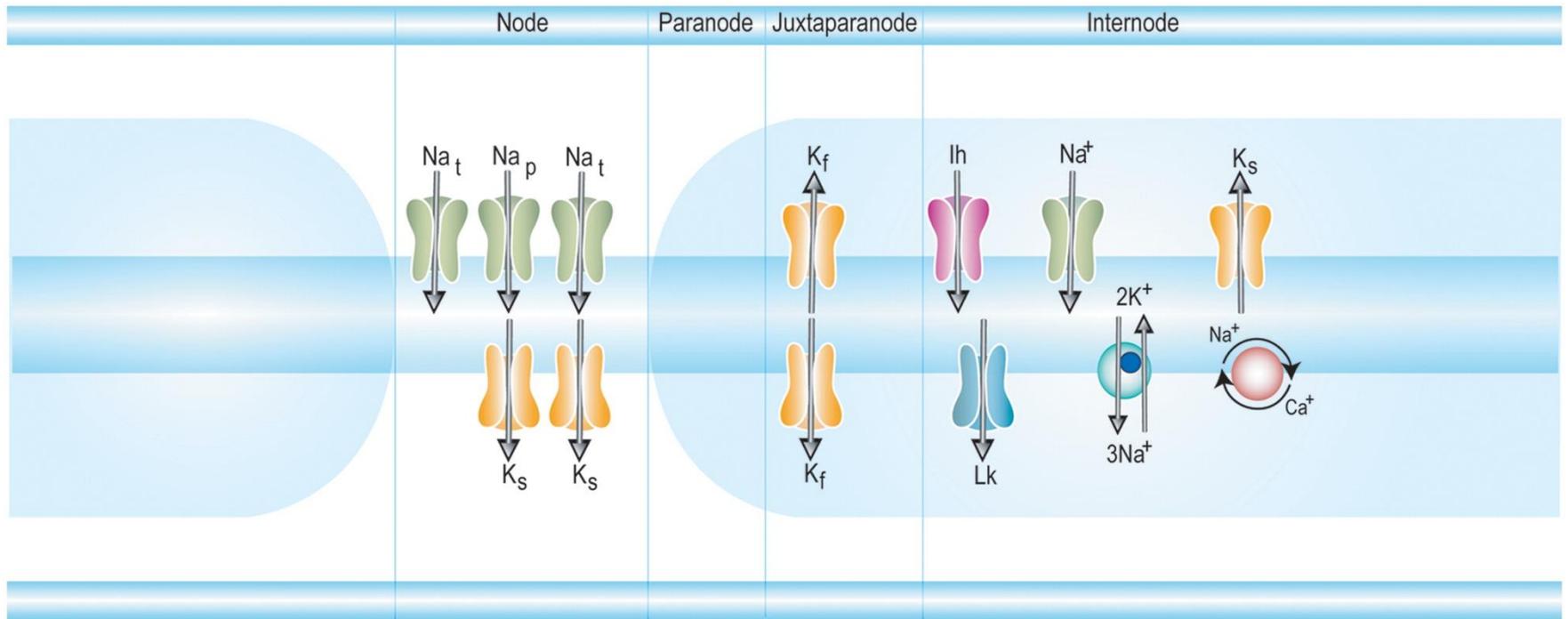
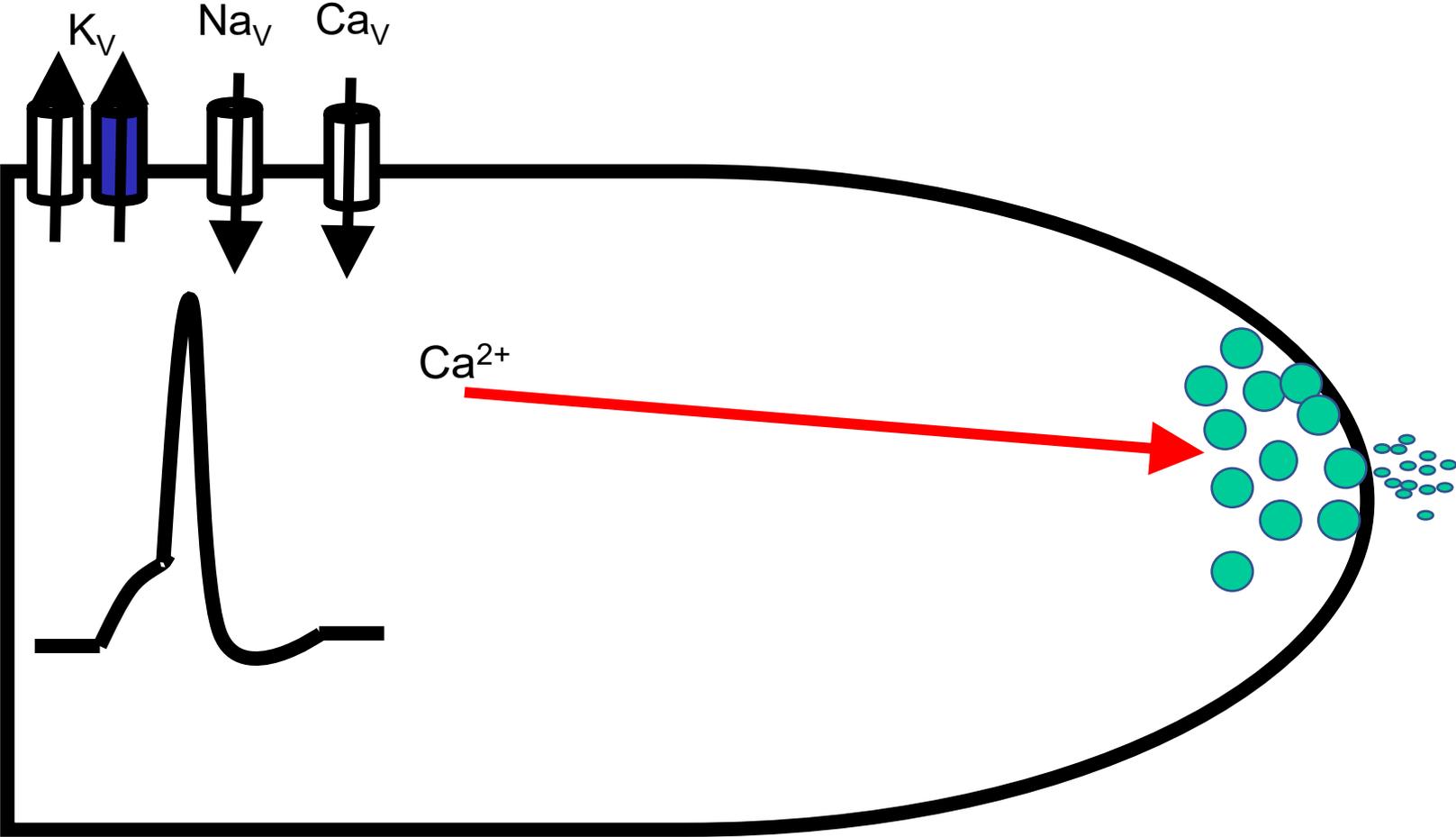


Figure 1. An overview of the various ion channels, pumps and exchangers that are important in axonal impulse conduction.  $Na^+$  channels are present in highest density at the Node of Ranvier as are slow  $K^+$  channels ( $K_s$ ). Fast  $K^+$  channels, also known as  $Kv$  1.1 channels are present in the highest concentration in the juxtaparanodal region and are blocked by 4-aminopyridine (fampridine). Inwardly rectifying channels ( $I_h$ ) are permeable to  $Na^+$  and  $K^+$  ions and limit axonal hyperpolarisation. Transient  $Na^+$  channels ( $Na_t$ ) are responsible for action potential propagation while persistent  $Na^+$  conductances ( $Na_p$ ) modulate resting membrane potential. The sodium-calcium exchanger ( $Na^+/Ca^{2+}$ ) plays an important role in the processes of axonal degeneration.  $Lk$  refers to leak conductances.

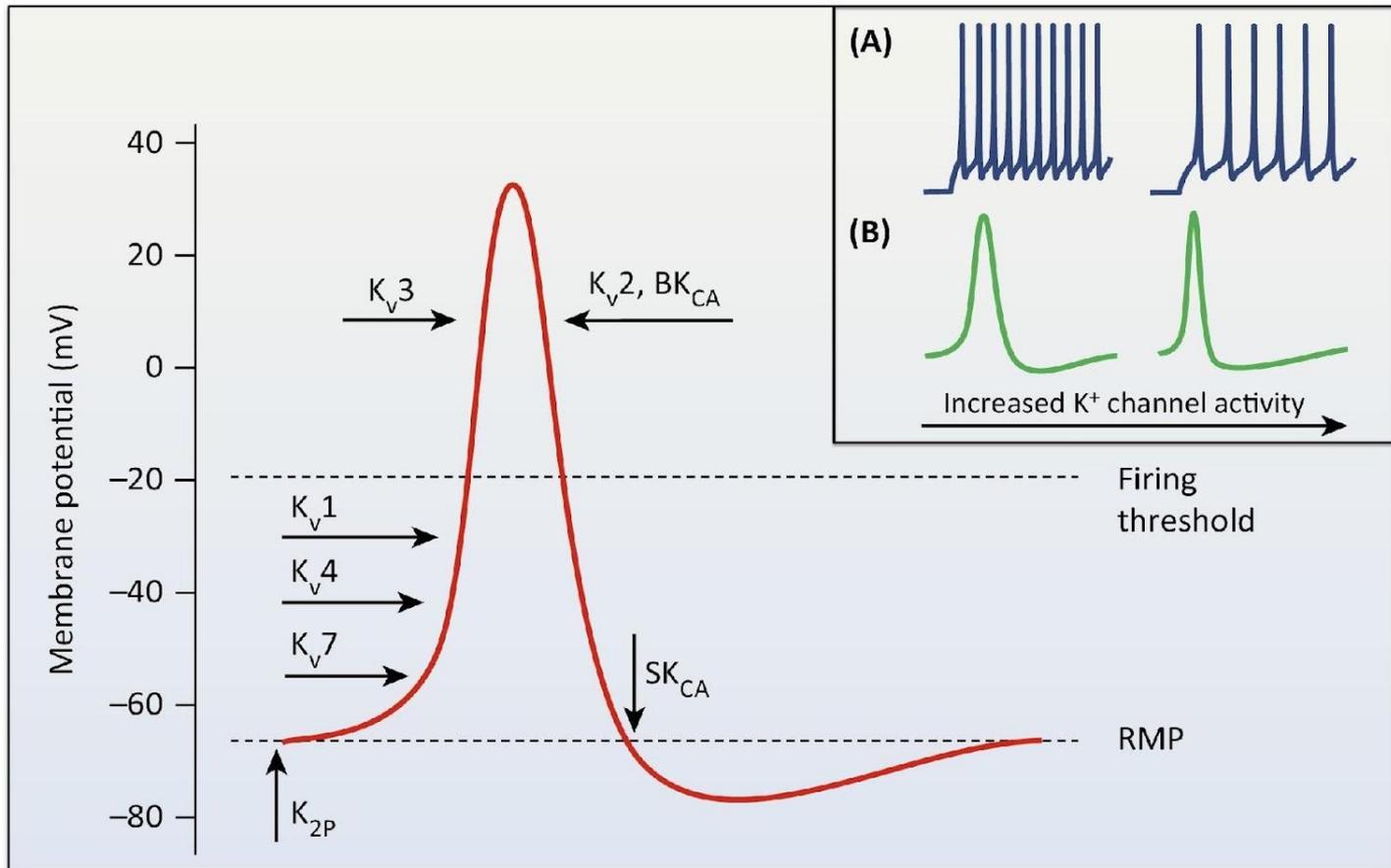
4-AP sensitive one



## Goldman-Hodgkin-Katz (G-H-K) equation

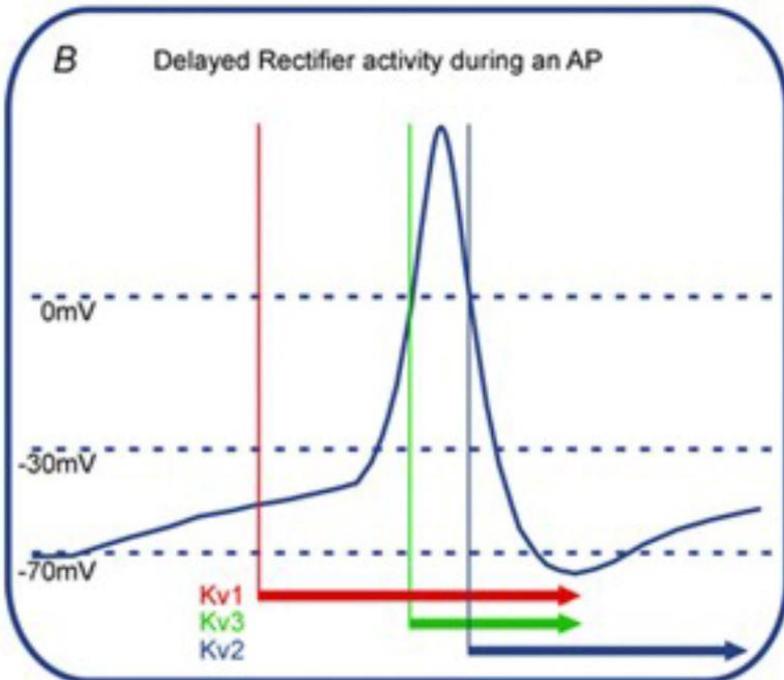
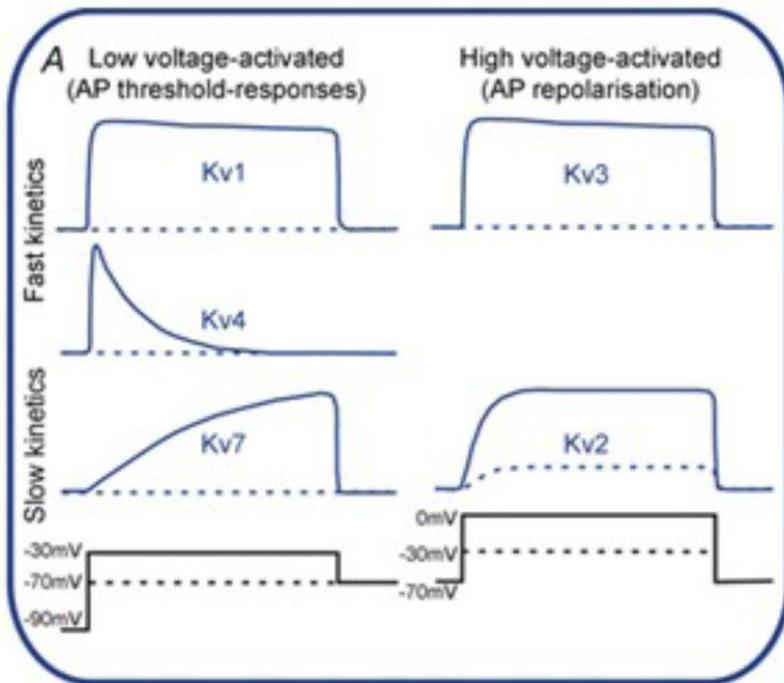
Here is a generalized G-H-K equation for Na<sup>+</sup>, K<sup>+</sup>, and Cl<sup>-</sup> ions:

$$E_{m_{K,Na,Cl}} = \frac{RT}{F} \ln \ln \frac{P_{Na^+}[Na^+]_{out} + P_{K^+}[K^+]_{out} + P_{Cl^-}[Cl^-]_{in}}{P_{Na^+}[Na^+]_{in} + P_{K^+}[K^+]_{in} + P_{Cl^-}[Cl^-]_{out}}$$

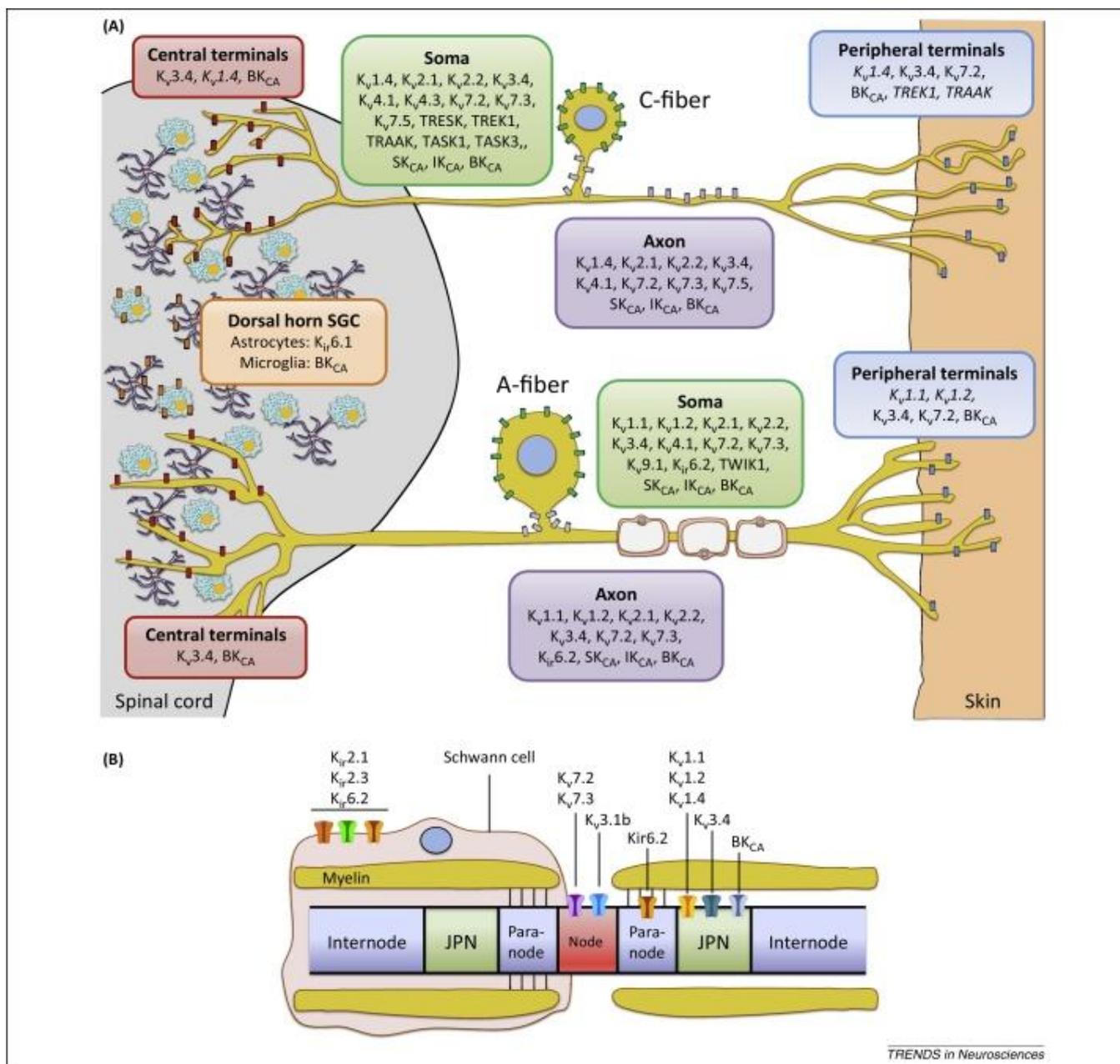


*TRENDS in Neurosciences*

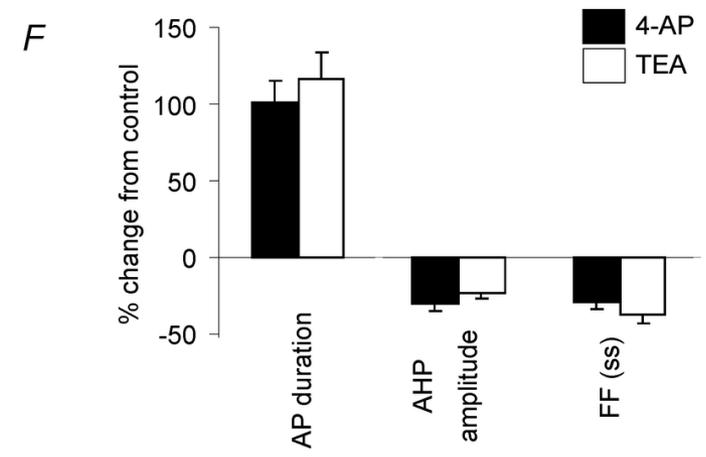
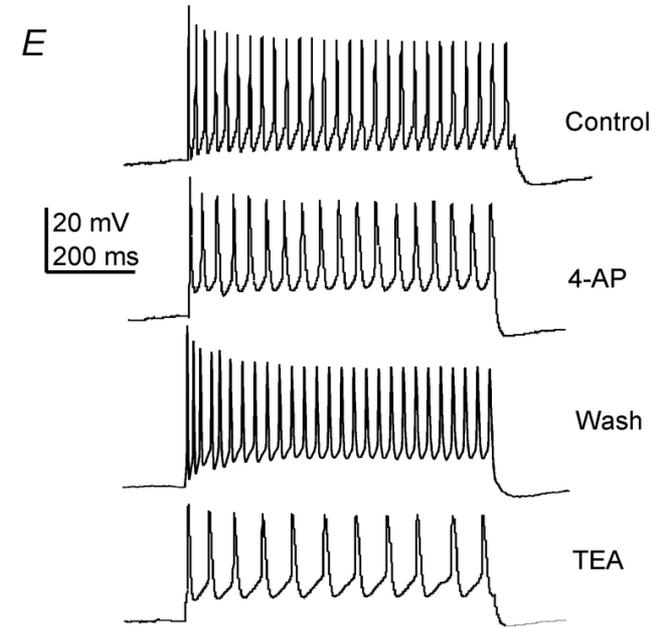
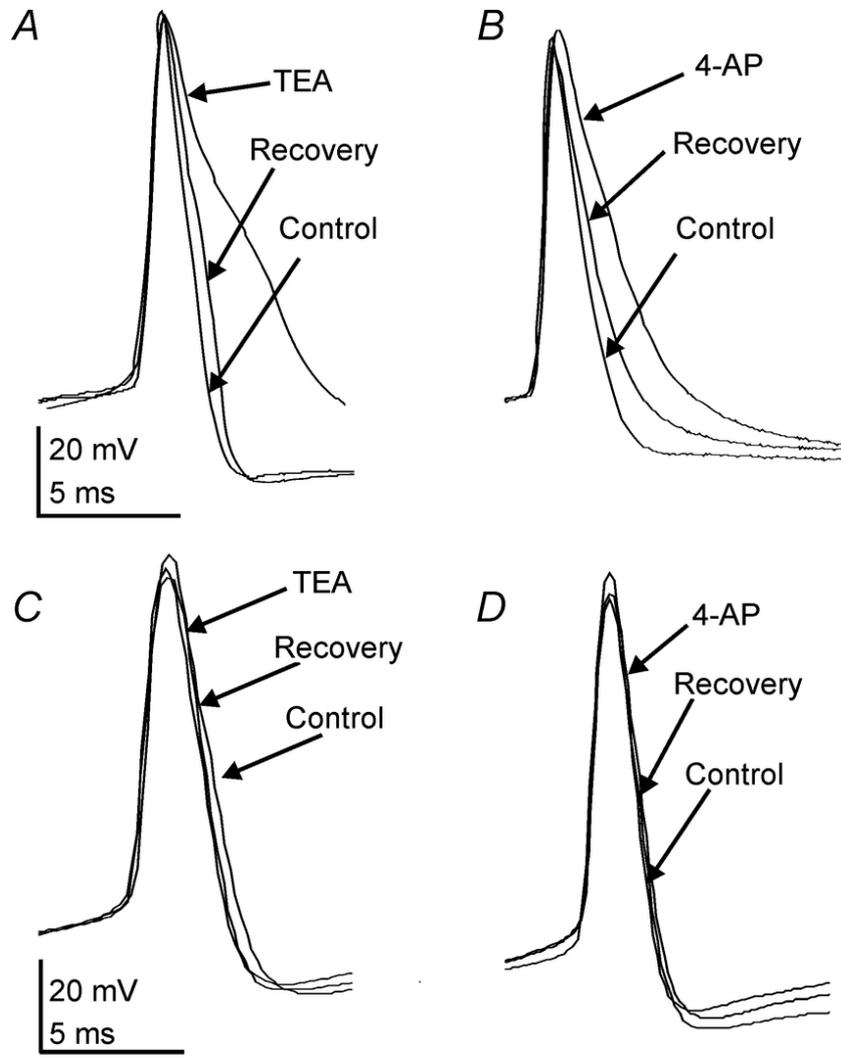
Tsantoulas C, McMahon SB. Opening paths to novel analgesics: the role of potassium channels in chronic pain. *Trends Neurosci.* 2014 Mar;37(3):146-58. doi: 10.1016/j.tins.2013.12.002. Epub 2014 Jan 21. PMID: 24461875; PMCID: PMC3945816.

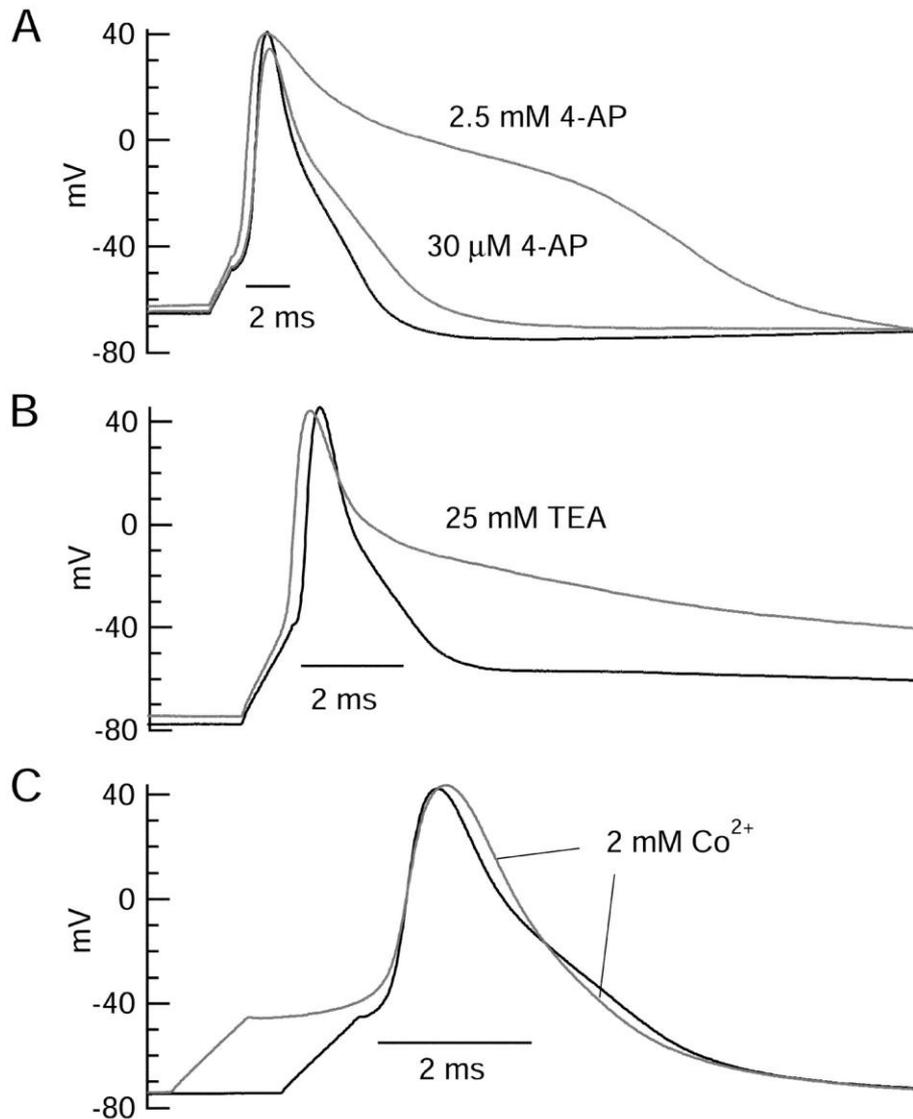


Johnston, J., I.D. Forsythe, and C. Kopp-Scheinflug, Going native: voltage-gated potassium channels controlling neuronal excitability. *J Physiol*, 2010. 588(Pt 17): p. 3187-200



Tsantoulas C, McMahon SB. Opening paths to novel analgesics: the role of potassium channels in chronic pain. *Trends Neurosci.* 2014 Mar;37(3):146-58. doi: 10.1016/j.tins.2013.12.002. Epub 2014 Jan 21. PMID: 24461875; PMCID: PMC3945816.





The overall voltage-dependent potassium current in the neurons could be split into three major components based on pharmacology and kinetics during step voltage pulses:  $I_D$  (fast activating, slowly inactivating, and sensitive to 4-aminopyridine at 30  $\mu\text{M}$ ),  $I_A$  (fast activating, fast inactivating, and sensitive to 4-aminopyridine at 3 mM), and  $I_K$  (slowly activating, noninactivating, and sensitive to external TEA at 3–25 mM). The potassium current during the action potential was composed of approximately equal contributions of  $I_D$  and  $I_A$ , with a negligible contribution of  $I_K$ .

<https://www.jneurosci.org/content/22/23/10106>