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## Characterizing the transient K<sup>+</sup> current contribution to subthreshold membrane potential oscillations in a hippocampal interneuron model

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A class of hippocampal interneurons in the CA1 region, bordering the lacunosum moleculare and radiatum hippocampal layers (the LM/R cell), has been shown to exhibit membrane potential oscillations (MPOs) subthreshold to action potential generation [1,2]. These oscillations occur at theta frequency (4–12 Hz) and are of interest because of their putative role in promoting theta activity at the network level. MPOs can be generated in the absence of synaptic input suggesting that they originate through interactions in the cell's repertoire of currents. In particular, the transient A-type K+ current ( $I_A$ ) has been shown to contribute [2]. In this study we present a model of the LM/R cell and show that  $I_A$  can enhance MPOs through modulation of the action potential threshold.

A single compartment model of the LM/R cell is developed using experimental data. I<sub>A</sub> is based on the Kv4.3 model given in [3], and the remaining currents, which consist of fast and slow delayed rectifiers, transient and persistent sodium currents, and a leak current, are modeled as Hodgkin and Huxley type currents. We also include applied current and white noise terms. The noise is modeled as either voltage independent or proportional to current magnitudes.

We consider the behaviour of the model system at voltages hyperpolarized relative to action potential threshold.

We define the subthreshold response function of the system for a given voltage and frequency as the complex ratio of voltage to total current for the currents driven by a small amplitude sinusoidal voltage around the fixed point. Here the voltage is a fixed function of time (as in idealized voltage clamp) and the currents follow passively. This ratio is preserved in the non-driven case in the vicinity of the fixed point and we can use it to predict the voltage response of the system to a given input signal. Simulations agree well with the theory and indicate that the assumed linearity is valid for physiological parameter ranges.

We assess the role of  $I_A$  by computing the difference between subthreshold response functions obtained with and without  $I_A$ . Blocking  $I_A$  can result in a hyperpolarizing shift in threshold depending on the  $V_{1/2}$  value of  $I_A$  inactivation. For noise independent of voltage,  $I_A$  promotes subthreshold power at theta for larger values of the shift while for noise dependent on current activation, the presence of  $I_A$  results in a more consistent enhancement of subthreshold theta power.

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