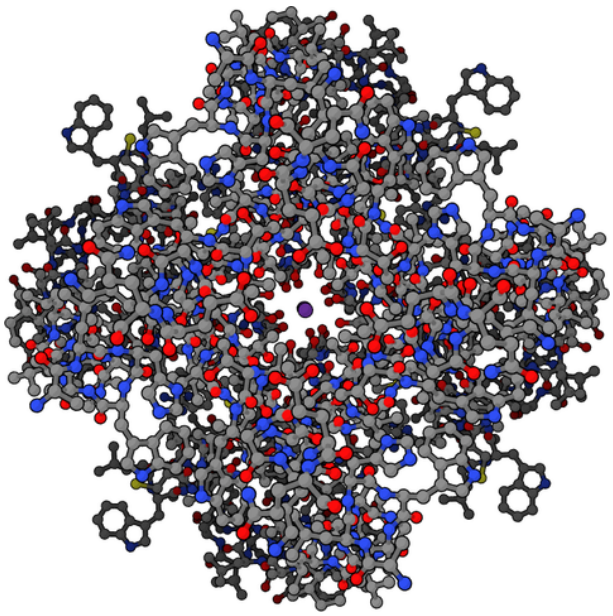

Figure 1: An open potassium channel, picture from wikipedia, which in turn took it from the Protein Data Bank. `http://en.wikipedia.org/wiki/Potassium_channel`

Gated channels

The nonlinear dynamics that neurons rely on to form spikes arise from the **voltage-gated channels**; these are ion channels whose conductance varies as the voltage varies. They are tiny molecular machines, crucially they are ion selective, only sodium ions can pass through a sodium gate, only potassium ions through a potassium gate. Each individual gate has a number of different gating states, we will briefly examine this, but ultimately each one is either open or closed, the overall smooth variation, though rapid, variation in these conductances comes from average a large number of individual discrete step-like changes as the individual gates open and close.

The potassium channel is a **persistent** gate; this is actually a little complicated, but roughly speaking it has one type of closed state and one type of open state; the sodium channel, which we will look at after the potassium channel also has one open state, but it has two types of closed states.

The potassium gate is actually composed of four independent subgates, all these gates must be open to allow potassium ions through, but each has its own dynamics. The membrane is usually modelled as having overall potassium conductance

$$g_K = \bar{g}_K n^4 \tag{1}$$

where $\bar{g}_K$ would be conductance if all the channels were open and n is the probability an individual subgate is open so n^4 is the probability an individual gated channel is open. The dynamical equation for n is quite complicated, it is of the standard form

$$\tau_n(V)\frac{dn(t)}{dt} = n_\infty(V) - n(t) \tag{2}$$

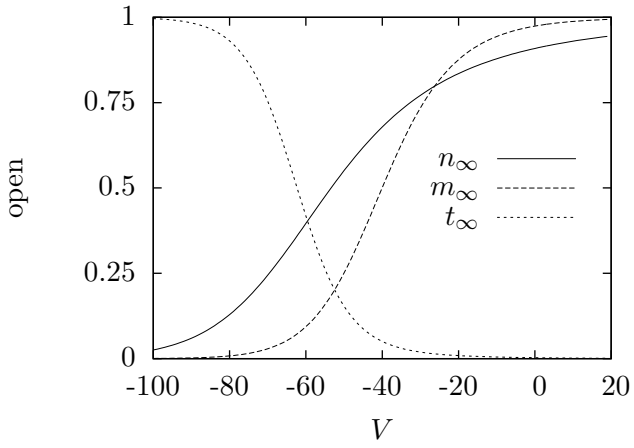

Figure 2: The asymptotic values of the gating probabilities.

If $\tau_n(V)$ and $n_\infty(V)$ where constant this would be simple, $n(t)$ would decay to n_∞ with a timescale of τ_n, however they aren't constants, they are functions of the membrane potential.

A graph of $n_\infty(V)$ is shown in Figure 2. We can see that n is small when the voltage is near the resting value but climbs towards one as V increases. Now, n isn't equal to n_∞, rather it decays towards it with a time constant given by $\tau_n(V)$, but we can see that the potassium channels open as the voltage increases. Before looking at how these play a role in spiking we will look at the sodium gates. It is worth noting though the way having four independent gates makes the dynamics crisper: if n is near zero, n^4 even nearer to zero, if n is close to one, n^4 is even closer. We also need to discuss reversal potentials. You would expect the flow of potassium to be dermined by $g_K V$, but it isn't; because there are more potassiums inside the cell than outside they would flow out even if $V = 0$. In fact we assume this doesn't change Ohm's law, the relationship between potential difference and current, rather, it just changes the zero point:

$$I_K = g_K(V - E_K) \tag{3}$$

where $E_K = -70$ mV, approximately, is called the reversal potential and can be calculated using an equation called the Nernst equation.

The sodium channel is called a transient channel because it has two closed states and one open one; generally its dynamics during the spike is

$$\text{closed I} \rightarrow \text{open} \rightarrow \text{closed II} \tag{4}$$

After that there is a slower process of resetting. The part of the gate that is closed to give the initial closed state is very like the sodium gate, but with three subgates; the probability of these subgates being open is usually called m; the other part, the gate that closes to give the second closed state is different in that it is not made of subgates, its pobability of being open is usually called h and its asymptotic value, h_∞, is near one for lower V and near zero for larger. Figure 2 includes graphs of $m_\infty(V)$ and $h_\infty(V)$. Finally, the reversal potential for

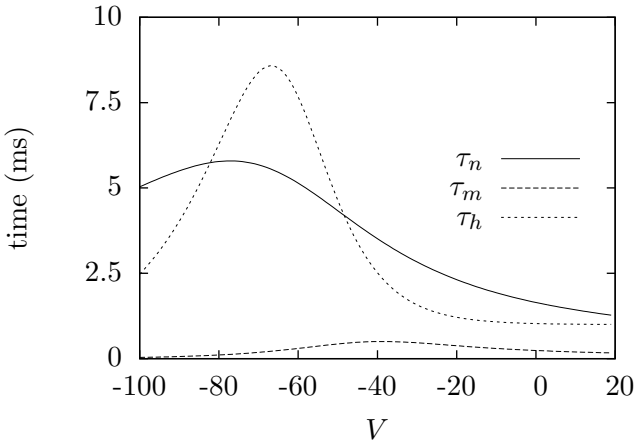

Figure 3: The time constants for the gating probabilities.

sodium is $E_{Na} = 50$ mV. The sodium current is therefore

$$I_{Na} = g_{Na}(V - E_{Na}) \tag{5}$$

with

$$g_{Na} = \bar{g}_{Na} m^3 h \tag{6}$$

We can now give a rough description of how spikes are formed. The time constants τ for the three gating probabilities are given in Figure 3, these are quite complicated, but the key thing is that τ_m is very small, no matter what the value of V is. This means that m stays very close to its asymptotic value m_∞. As V approaches the threshold of about -55 mV, m increases towards one, with m^3 increasing even more dramatically. Opening the sodium gates allows sodium to flood the cell, increasing the V further and further opening the gates. This gives the rapid upswing in voltage, the rising part of the spike. The other two gating probabilities have slower dynamics and it takes n and h a while to catch up with n_∞ and h_∞. However, as h decreases, it closes the sodium gates again, preventing more sodium getting in to the cell; n increases opens the potassium gates, potassium flows out reducing the V again, back towards -70 mV. This gives the downswing of the spike. Afterwards everything resets.

All of this together gives the Hodgkin-Huxley equation, basically it equates the rate of change of V to a set of currents, the leak current giving the roughly linear behavior below threshold we saw in the integrate and fire model and the gated channels forming the spike.

$$C_m \frac{dV}{dt} = \text{currents} \tag{7}$$

For a more accurate model further channels, and therefore further currents, can be added, other sodium and potassium channels with different dynamics, or a calcium channel. It is also common to investigate models 'between' the integrate and fire model and the Hodgkin-Huxley equation which add some of the nonlinearity to the integrate and fire dynamics.