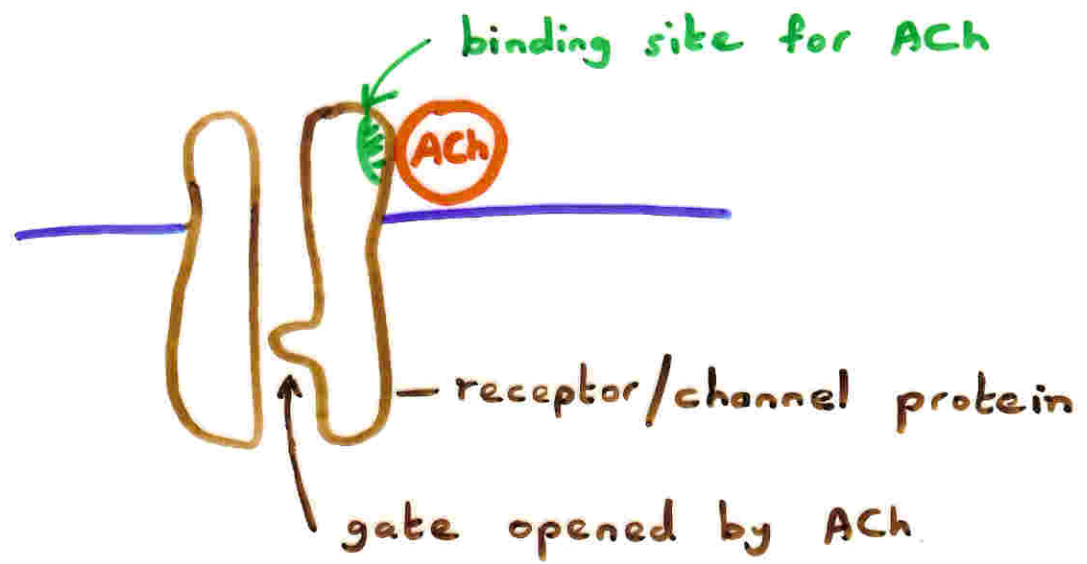


# Neurotransmitter Receptors

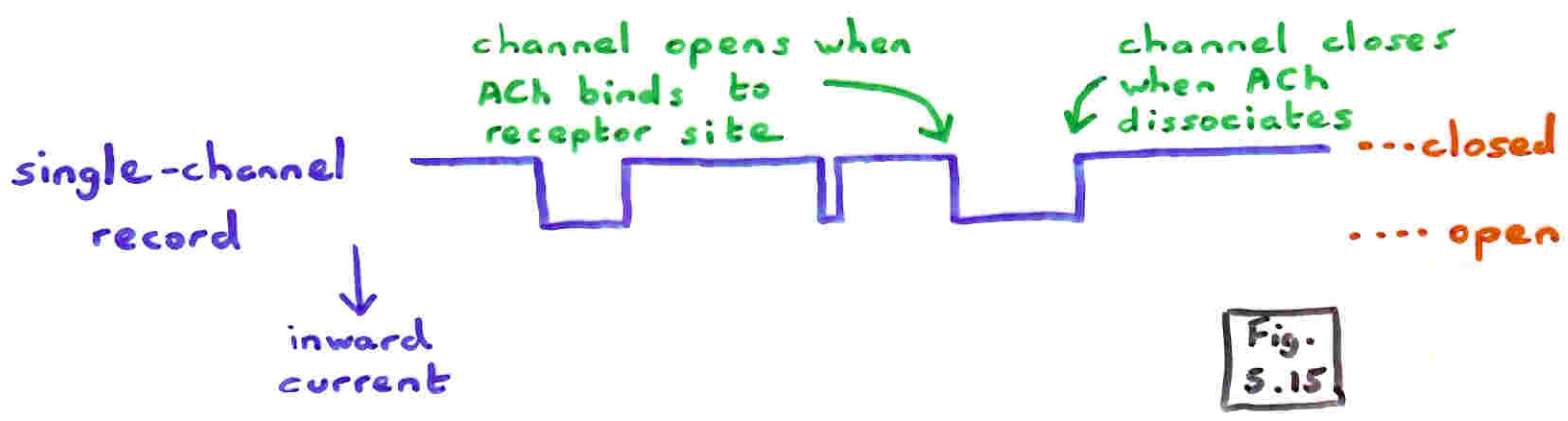
## End-plate channels



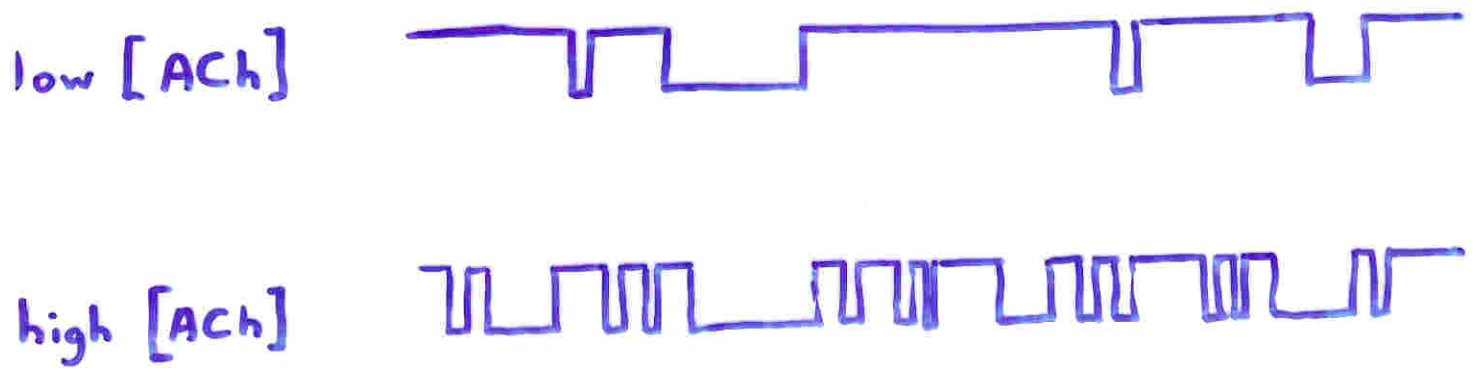
endplate channel is integral protein (5-subunits) forming ion channel and receptor for ACh.

Channel is permeable to both  $Na^+$  and  $k^+$

# Channel gating



increasing  $[ACh]$  increases frequency of channel openings; not duration of opening or size of current through channel.



Overall current at endplate increases with increasing  $[ACh]$  because more channels are open at any instant of time.

## Channel opening requires binding of 2 ACh Molecules

ACh-activated channel formed from 5 subunits

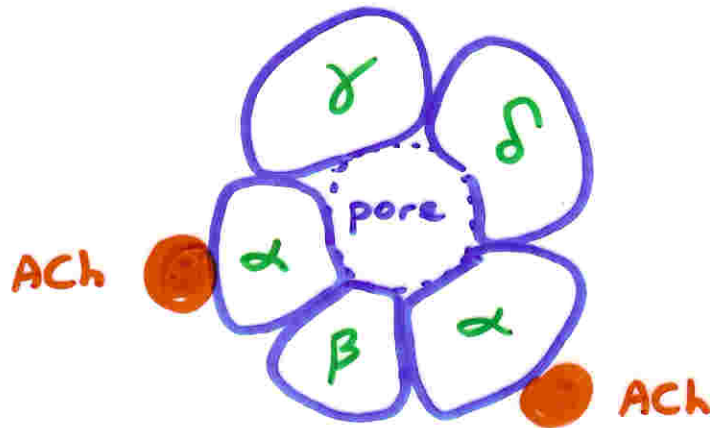


Fig. 6.3B

ACh-binding sites are on  $\alpha$  subunits. There are 2 of these in each channel.

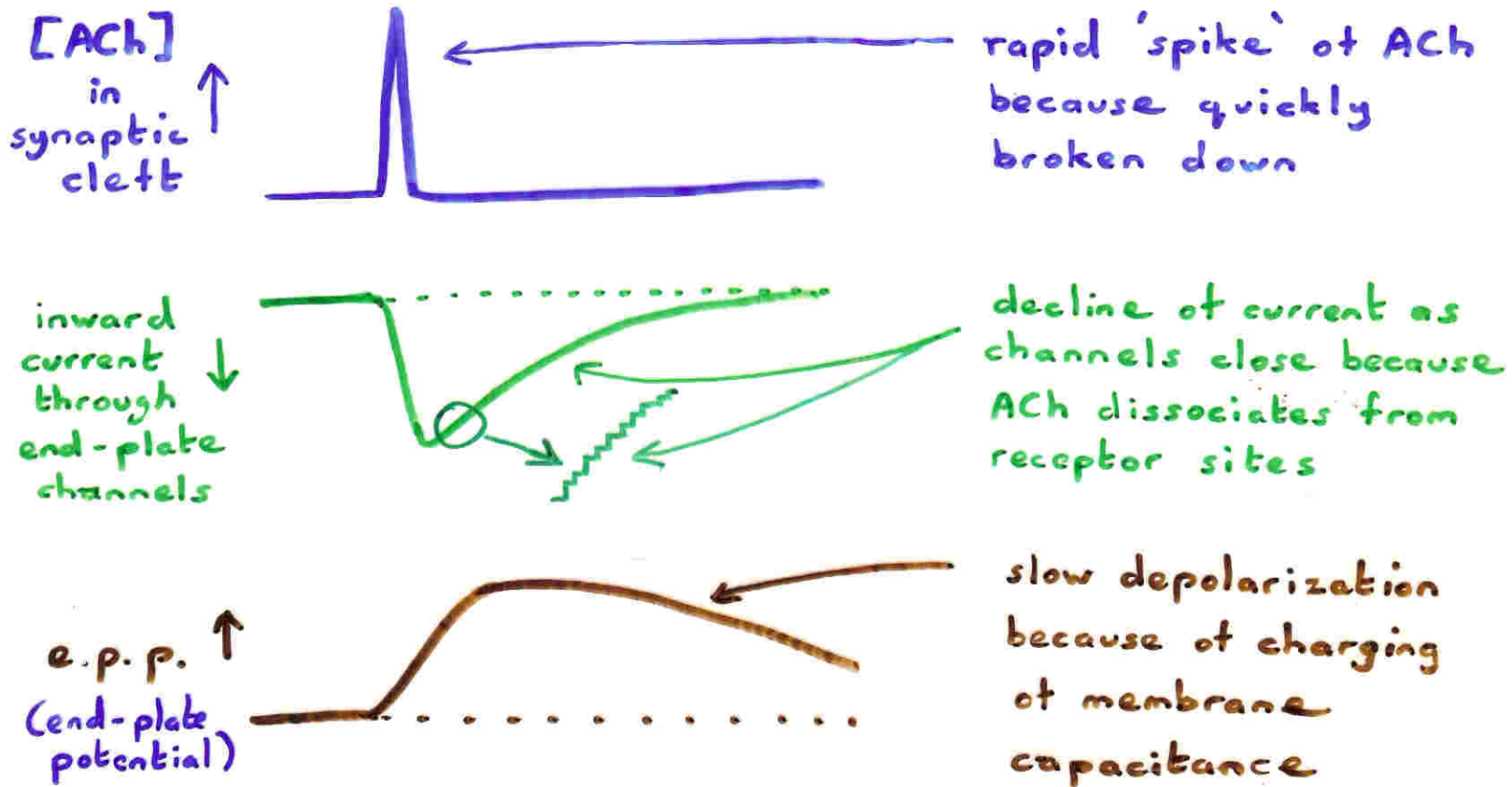
ACh must bind to both sites before channel opens. Thus, channel opening increases as the square of  $[ACh]$ .

[eg. if double  $[ACh]$ , probability that a single site will bind ACh doubles.

Thus, probability that both sites bind ACh quadruples]

# Time course of channel openings during e.p.p.

ACh released into synaptic cleft is very quickly broken down by enzyme, acetylcholinesterase. [hydrolyzed to choline and acetic acid : choline is recycled by nerve to make more ACh]



## Ionic basis of the end-plate potential

End-plate channels let through  $\text{Na}^+$  and  $\text{K}^+$  ions. Equilibrium potential is thus midway between equilibrium potentials for  $\text{Na}^+$  (+60mV) and  $\text{K}^+$  (-90mV).

Voltage-clamp muscle and record end-plate current that flows when clamp at different potentials.

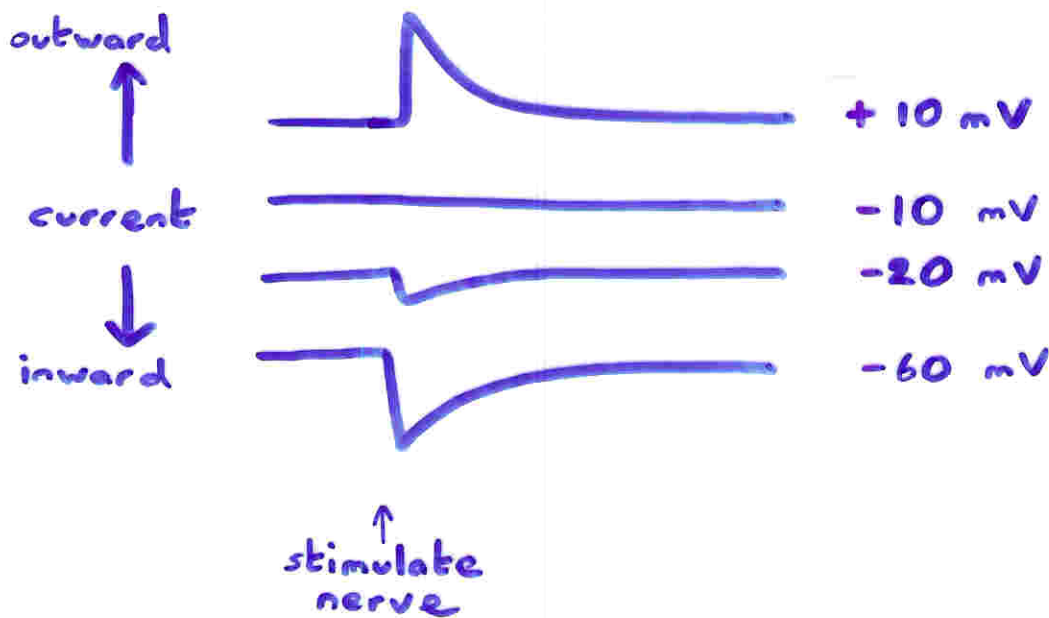
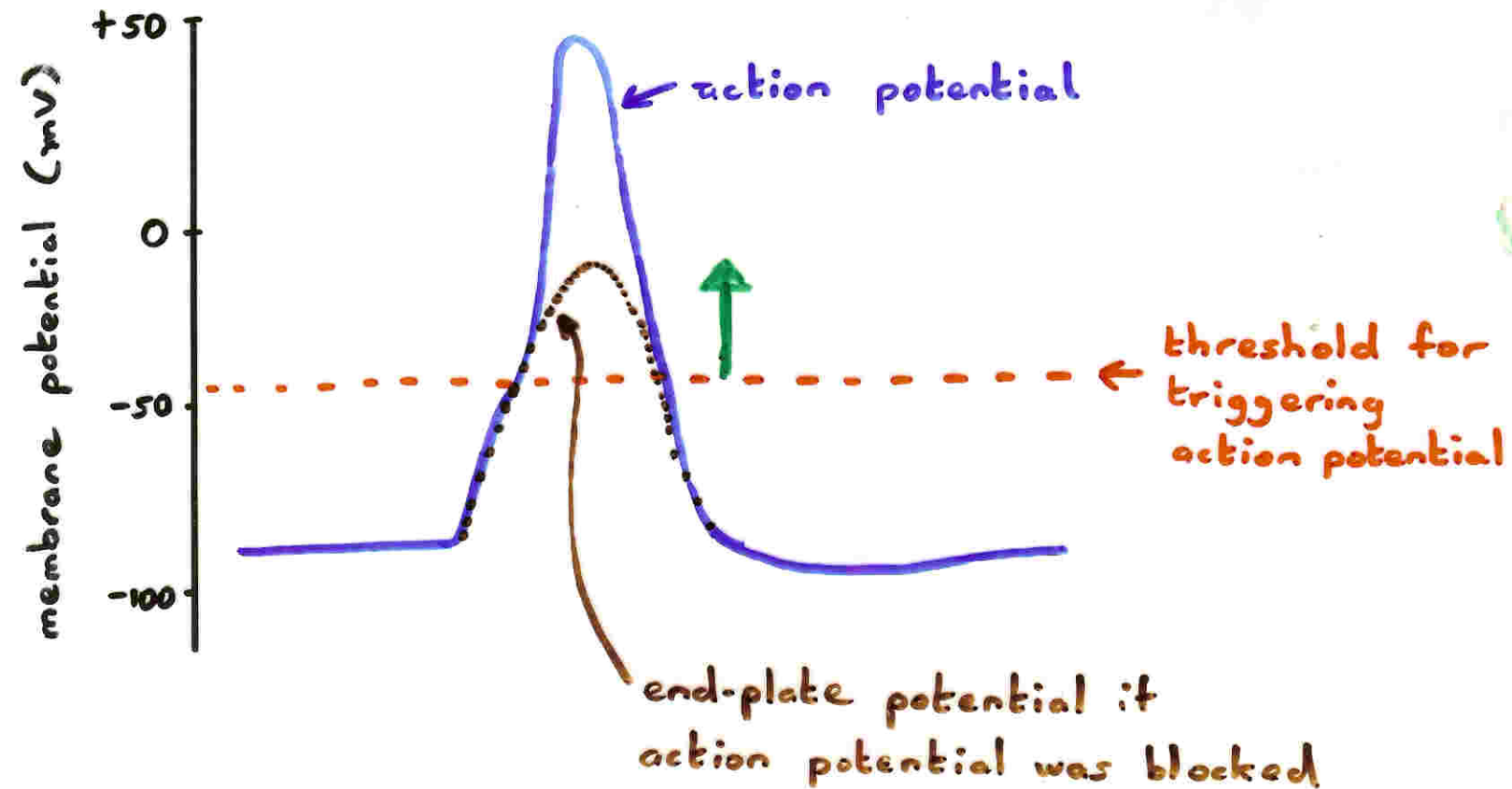


Fig. 5.16

No matter how much ACh is released, the end-plate potential cannot depolarize a muscle fibre beyond -10mV.

However - depolarization to  $-10\text{mV}$  is more than enough to trigger an action potential.



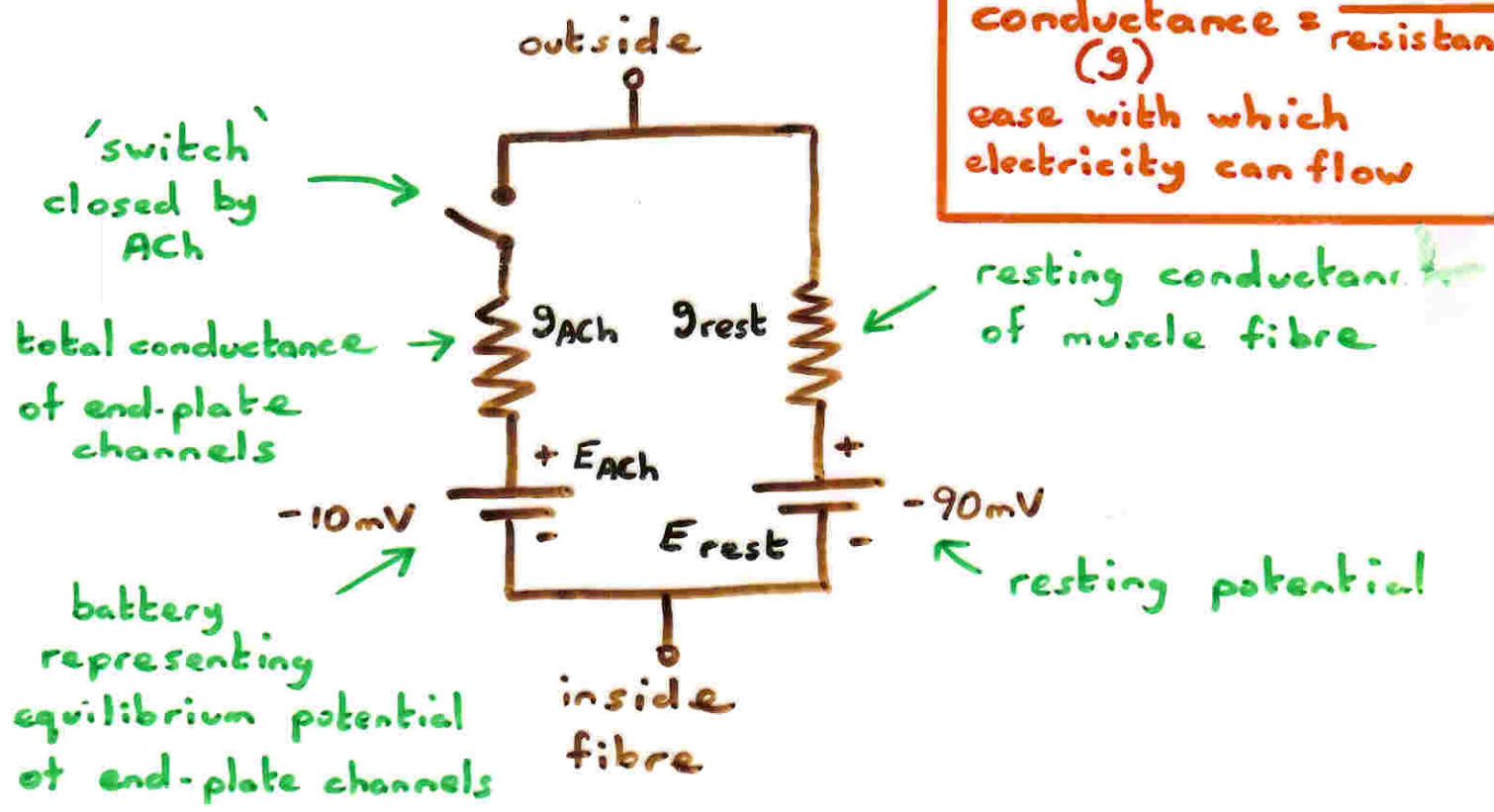
# Equivalent electrical circuit for endplate

Resting potential tends to hold muscle fibre at -90mV.

Current through end-plate channels tends to depolarize to -10mV.

Resulting potential depends on relative conductance of end-plate (how many channels open)

conductance =  $\frac{1}{\text{resistance}}$  (g)  
ease with which electricity can flow



final voltage of muscle  $E = \frac{g_{ACh} \times E_{ACh} + g_{rest} \times E_{rest}}{g_{ACh} + g_{rest}}$