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Membrane biophysics (Sargent)

Neurophysiology Primer

- Properties of resistors, capacitors in series/in parallel
- Kirchoff's current law/Kirchoff's voltge law/Ohm's law
- Properties of RC circuits
 - Biological membranes as parallel RC elements
 - Neuronal geometry and the passive electrical properties of excitable cells

Determinants of Membrane Potential

- Model cells permeable to one ion
- The Nernst Equation equilibrium
- Model cells permeable to more than one ion steady state
- Current, conductance, and driving force (Ohm's law)
- The Goldman/GHK equation
- Contribution of pumps to membrane potential
- Leak channels and the resting potential

Action Potentials

- Voltage clamp; HH analysis
- Modeling
- IV curves
- Threshold
- Refractory period
- Gating current
- Propagating action potentials

Single Channels

- Noise
- Patch Clamp
- Popen
- Two state transitions; deriving rate constants from dwell times

Ion Channel Structure and Function (Kirichok, Jan L)

Ion channel classes by selectivity

K channels

- K_v channel structure
 - Selectivity filter
 - Gating
 - Voltage sensor
 - Inactivation
- Kir channels
 - Structure
- Na channels
 - Na_v channels

Ca channels

Ca_v channels

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Selectivity Auxiliary subunits Non-selective cation channels CNG channels, HCN channels Nicotinic acetylcholine receptor Structure Permeation Gating Synaptic Transmission (Sargent) **Electrical Synapses Ionotropic Receptors** Role of ionotropic receptors in fast chemical transmission Determinants of the time course of synaptic current Ionic basis of excitatory and inhibitory synaptic transmission **Reversal potential** Inhibition by shunting **Transmitter Release** Role of calcium Fate of released transmitter Direct and indirect methods for measuring transmitter release Spontaneous and evoked transmitter release Synchronous and asynchronous release The quantal hypothesis The vesicle hypothesis Statistical methods Use of binomial/poisson statistics to measure changes in p_v Estimating N, P, and Q from variance-mean analysis Univesicular and multivesicular release *Linearity/nonlinearity of synapses* Short-term Facilitation/depression Glutamate Receptors and Plasticity (Nicoll) Glutamate Receptors (Nicoll) Classification **Expression cloning** The postsynaptic density (PSD) **AMPARs** Structure Gatina Kinetics and subunit composition Genetic dissection of AMPAR subunits AMPAR trafficking **NMDARs** Subunits and properties Developmental switch in subunits

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Scaffolds Adhesion Proteins Plasticity (Nicoll) Taxonomy of memory systems **Hippocampal LTP** Induction of LTP/LTD Associativity AMPARs, NMDARs Expressions Mechanisms of LTP/LTD "Quantal Analysis" and LTP Silent Synapses CaMKII Receptor Centric Model vs PSD centric Model Integration (Bender) Cable theory Role of capacitance in filtering Neuronal exploitation of passive cable properties Fast integration Time difference coding Sublinear integration Passive integration in dendrites Orthodromic and antidromic spike propagation Active properties of dendrites Importance of synaptic location, EPSPS vs IPSPs Synapses onto spines and shafts; what are spines good for? Axonal integration Analog vs. digital components of action potential induced release Determinants of AP initiation Modification of integration Neurotransmitter transport (Edwards) Synaptic Vesicles Purification Proteins associated with synaptic vesicles Transport of synaptic vesicles Molecular basis of synaptic vesicle docking and priming Molecular basis of calcium-dependency of exocytosis Molecular constituents of the active zone Peptidergic vesicles Neurotransmitter transport Endocytosis Clathrin **Adapters** BAR domain proteins

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Uncoating

Models for exocytosis/encocytosis

Synaptic vesicle pools

Are spontaneous and evoked vesicles different

Receptor Pharmacology (Whistler)

Ionotropic and metabotropic receptors

G protein mediated signaling

Primer on pharmacological terms (ligand, agonist, antagonist)

Ligand affinity

Efficacy

Ligand potency and selectivity

Allosteric modulators

Receptor oligomerization

Receptor desensitization/downregulation

Biased agonism