

Figure 6-1. Organization of skeletal muscle, from the gross to the molecular level. F, G, H, and I are cross sections at the levels indicated in the levels in the levels in the levels indicated in the levels in



General Physiology
Second Semester 2023
Lecture 14 and lecture 15
Organization of the Nervous System,
Neuronal Graded Potentials
Basic Functions of Synapses, and Synaptic Potential (EPSP and IPSP)
Neurotransmitters

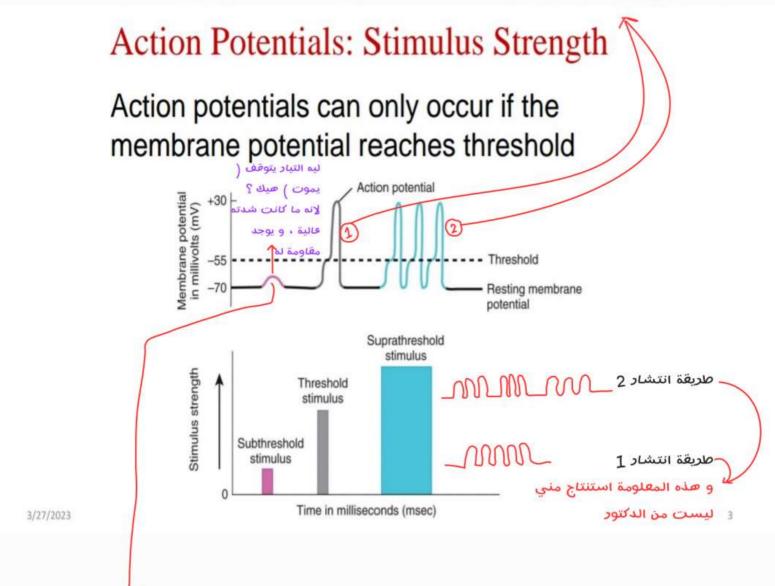
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Lectures objectives

- Define neuronal graded potentials and list examples of graded potential and their functional significance
- Describe the main morphologic features of synapses.
- Distinguish between chemical and electrical transmissions at synapses.
- · Describe steps during chemical synaptic transmission
- Describe excitatory and inhibitory postsynaptic potentials, outline the ionic fluxes that underlie them, and explain how the potentials interact to generate action potentials
- List the primary neurotransmitters in the nervous system (Acetylcholine, biogenic amines and amino acid neurotransmitters and nitric oxide
- Identify different peptides neurotransmitters

3/27/2023

هسا هذا all-or-none و هو من خصائصه انه potentials لكن مش كد ال potentials كلك، في action potential كلك، في و عد ال graded potential بكون مش action potential و هو الا action potential بكون مش



هون صار تحفيذ نتج فنه depolarization، لكن هذا التحفيذ فير كافي ليصل إلى جهد الفتبة بالتالي لم يتم توليد action potential، بقي هذا التحفيذ في مكانه و لم ينتقل

بدي إياك تركز على فكرة انه الجهد الناتج عن المحفذ إلى رفع جهد الخلية إلى جهد العتبة هذا لن يكمل مسيرته على طول محود العصبونات ، هذا فقط يوصل جهد الخلية إلى العتبة و ايصاله هذا سوف يقوم بإنشاء action على طول محود العصبون ، يعني خاصية potential

Means that the amplitude depend on the strength or the amount of stimulus. In other way, if you have generate a localized potential and it is stimulated by neurotransmitters, the higher amount of neurotransmitters, the greater amount of amplitude as a result of the increase in neurotransmitters.

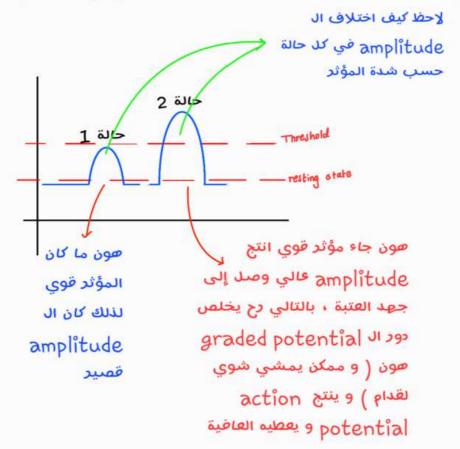
هذا موجود في بعض الأعصاب Graded Neuronal Potentials بيس كلها (لها تعريف في سلايد Small localized Changes (deviations)) in resting membrane potential initiated by an appropriate stimulus Can be depolarizing hyperpolarizing تم جلبها Graded potentials are brought about by external stimuli (in o sensory neurons and they are known as receptor or generator potential) or by neurotransmitters released in synapses, M where they cause graded potentials in the post-synaptic The changes in membrane potentials membrane is due to Time in milliseconds (m activation of different types of channels and changes in (a) Hyperpolarizing graded potential permeability of different ions like Na+, K+, or Cl-. هو سبب تواصل الخلايا في أطراف القصبونات ح (mV) in militoits (5 · Graded potentials are the only means of communication used by some neurons play very important roles in the initiation and integration of the long-distance signals by Time in milliseconds (msec) neurons ترجعة الإشارة و (b) Depolarizing graded potential لما يكون فندي إشارة بدى انقلها مسافة فهمها 3/27/2023 طويلة عبر الأعصاب رح تلاقي انه سبب انتقالها على طول المسافة هو تواصل العصبونات مع بعض عن طريق ال graded

(# localized change: potential don't travel to long distance, you can find them at a small distance from the site of stimulation

يعني ما رح تعمد self-propagation و توصد إلى terminal معين ، بس رح تعمد شوي حواليها ، و ذلك لأنه صدفها تعمد depolarization لجهد الخلية حتى يصد لجهد العتبة و من ثم عند الوصول لجهد العتبة رح يصيد عندي Action potential

بعد هذا الكلام الدكتور قال ؛ يعني ال localized potential سواء مباد في ال receptors او في ال action ومنان يصيد عندنا action مدخلة depolarization عشان يصيد عندنا synapses و ذلك مدخلية او نعمل action potential عشان نعمل stop عشان نعمل potential و ذلك neurotransmitters و خلك حسب نوع ال

هسا خليني اشرح لك الحكي المكتوب في السلايدين الي فوق ، هسا احنا عندنا شيء في الأعصاب اسمه graded potential محدد ، بل حسب قوة المؤثر الي اجاه و graded potential محدد ، بل حسب قوة المؤثر الي اجاه محدث ، بل حسب قوة المؤثر الي اجاه محدث ، و هذا له دور في تفعيل ال محدثان محدث ، لأنه إذا بتتذكروا حكينا انه عشان نولد محدثان محدث و هذا له دور في تفعيل الله و محدث الخلية لجهد العتبة لانه و مخد الخلية لجهد العتبة ، طيب لو تفكر فيها عشان يدفعه لجهد العتبة لانه عمون الدسمة الي رح ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي رح ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي رح ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي رح ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادسمها و متفيد حسب قوة المؤثر ، شوف الدسمة الي در ادبي القون الدين اله عدد العدد الع



هسا اول نقطة بدي إيالا تفهمها ، ثاني شفلة قالها الدكتود انه الا potential المهموم و المهموم ا

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expressed in dendrites and the soma, while β4 is primarily expressed in nerve terminals and the soma to a lesser extent (Wynne, Puig, Martin, & Treistman, 2009). BK channels in the presynaptic terminal are likely to provide local negative feedback of voltage-dependent Ca2+ influx during depolarization-secretion coupling, thus limiting secretion, while cell body BK channels may be important in regulating patterns of cell excitability (Dopico et al., 1999). BK channel potentiation by ethanol is decreased by chronic exposure and is thought to be an important factor in alcohol tolerance (Knott, Dopico, Dayanithi, Lemos, & Treistman, 2002). BK channel sensitivity to ethanol is splice variant dependent with BK channels expressing the ALCOREX insert (alcoholregulated exon) showing potentiation by ethanol (Pietrzykowski et al., 2008). The same study revealed that chronic alcohol exposure upregulates microRNA miR-9 which mediates the selective downregulation of BK channel a-subunits containing the ALCOREX exon, leading to alcohol tolerance.

The PVN also consists of a large population of parvocellular neurosecretory cells which project to the median eminence at the brain and release hypothalamic secretagogues, such as CRH, TRH, and AVP, into portal circulation to stimulate cells of the anterior pituitary (Kiss, 1988). In response to stress, CRH is released from the PVN, a small population of neuroendocrine cells comprising of approximately 4000 neurons in rate

Activated Potassium Channels

A. Alioua, ... L. Toro, in Reference Module in Biomedical Sciences, 2014

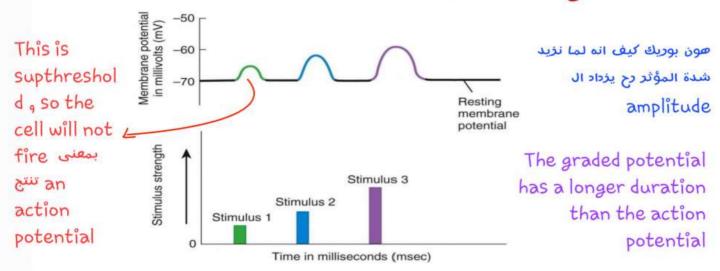
Abstract

The pore-forming a subunit of large conductance voltage-and Ca2+-activated K+ channels (KCa1.1) is widely expressed throughout the nervous system. Four K_{Ca}1.1 subunits are required for channel activity, which contributes to the control of neuronal pacing, transmitter release, and neurovascular coupling. In neurons, K_{Ca}1.1 channel activity can shape the repolarization and fast hyperpolarization phases of the action potential or control transmitter release via a negative feedback mechanism. K_{Ca}1.1-specific roles in each type of brain cells are beginning to emerge and depend on their geographic localization (e.g., soma, dendrites, pre-/postsynaptic terminals, plasma membrane, nucleus, mitochondria) and molecular constitution, which can vary depending on the K_{Ca}1.1 variants forming the channel, and also association or not with modulatory β_2 and β_4 subunits and/or other partner proteins. Silencing regulatory β_4 subunit expression causes temporal lobe seizures, whereas the absence of K_{C3}1.1 leads to deficit in locomotion and alteration of hearing and circadian firing rates. A K_{Ca}1.1 channelopathy relates to coexistent general FEEDBACK

طبعا لاحظ انه كلامنا في ال graded potential عن ال graded potential هذا يعني انه يحدث في ال chemical synapses، و تأكيد هذا الكلام انه حكينا في مسلايد 4 انه يحدث فقط في some neurons سلايد 4 انه يحدث فقط في some neurons التأكيد على هذا الكلام هو سلايد 13 الجدول ، شوف ال type of channels تاعت ال graded potential

هسا اذا حسيت حالك ما فهمت الشرح الي فوق عشان مومنوع ال neurotransmitters عادي ، كمل لآخر المحامنرة (سلايد 21) بعدين ارجع له

Graded Potentials: Stimulus Strength

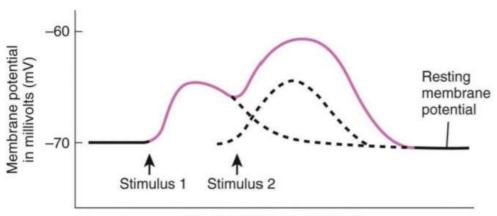


The amplitude of a graded potential increases by increasing stimulus intensity

3/27/2023

جمع (و هو معنى Graded Potentials: Summation

Graded potentials can be add together to become larger in amplitude



Time in milliseconds (msec)

مش احنا لما نحفذ عصبون بصيد graded potential، لكن لو مش كافي و ما وصل threshold تلاقيه طلع والمستعدة (صاد اله amplitude خفيفة) و بدجع بنذل ، لكن لو اتبعته ب stimulus آخد بسدعة قبل لا ينذل ال amplitude تاعه ، رح ينصاف هذا ال stimulus له و يدفع اله amplitude اكثد مساق الله عنه الله عنه الله المستعديد حوالي 10 حبات من الله و يدفع الله عنه كانوا كافيات و عنه بتحديد حوالي 10 حبات من الله و الله عنه الله الله عنه الله الله عنه و ملت stimulus آخد يحدد لك كمان neurotransmitters المستعدم وصلوا و ينضافوا هذول الله عنه الله عنه المستودة الله عنه و عملت وعملوا المستودة الله عنه و عملت neurotransmitters الجداد للي قدام و عملوا المستودة الله عنه الله ع

Graded potentials are localized and travel only a short distance form

sensory neurons الي حكينا عنه site of stimulation

الدكتور ما شدحها ، لكن رح اشرحها في سلايد ١٢ حسب فهمي

The membrane potential of a cell can be depolarized by using a stimulating current generator, and the potential can be recorded by a pair of electrodes, one inside the cell and the other in the extracellular fluid, (a)

Membrane potential is closer to the resting potential with increasing distance from the depolarization site.

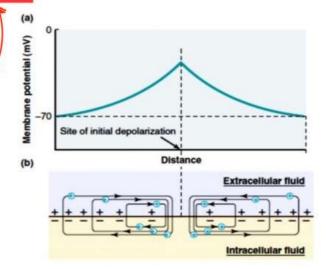
(b) Local current surrounding the depolarized region produces depolarization of adjacent regions.

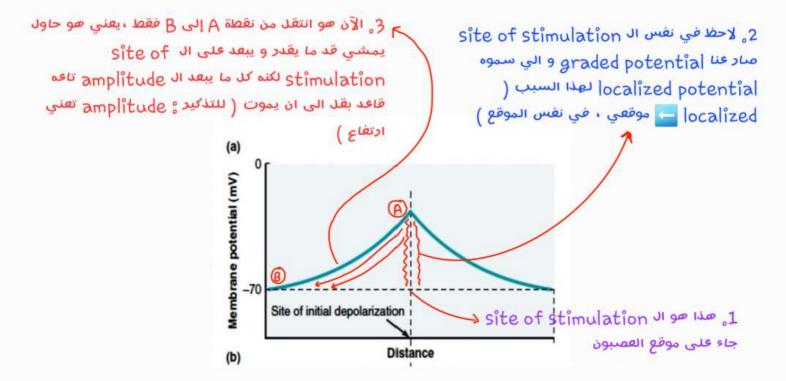
Because the electric signal decreases with distance, graded potentials (and the local current they generate) can function as signals only over very

short distances (a few millimeters).

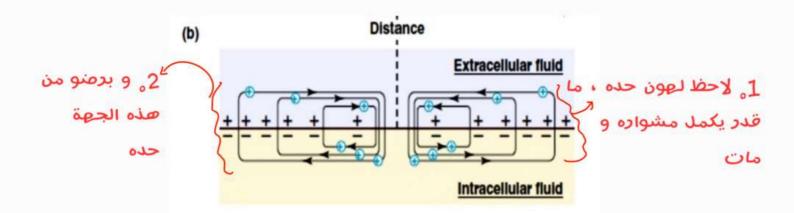
Thus, graded potentials <u>do not travel</u> along distance like an action potentials

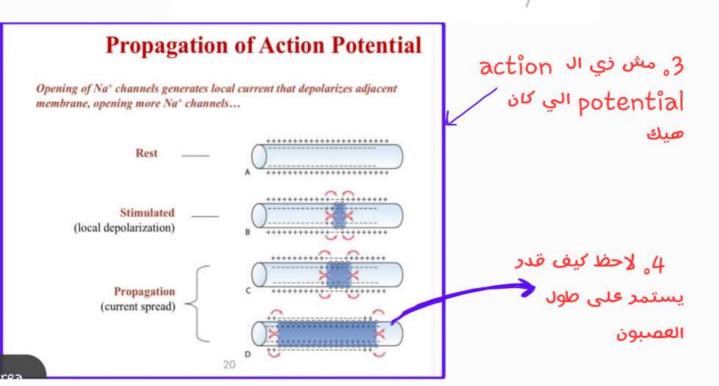
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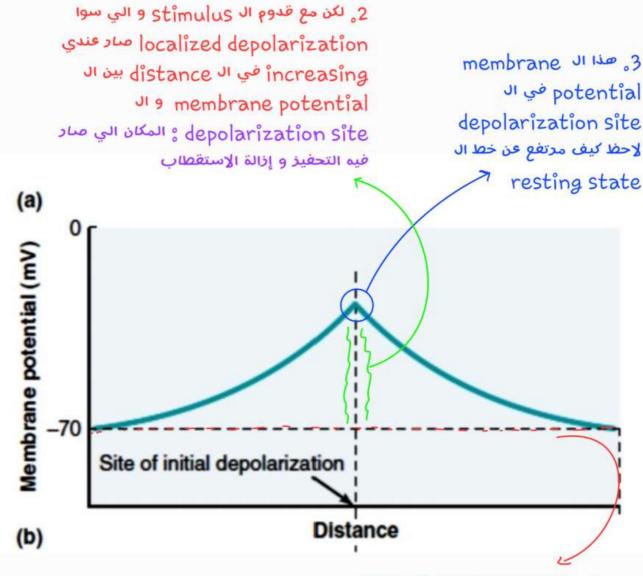


4. الآن هو في النهاية مات ، ما استمد على طول المحود ، يعني ما عنده self-propagration مثل ال self-propagration و مثل ال self-propagration و من ثم يموت او انه يوصل به محود اله potential و من ثم يموت او انه يوصل اله threshold و يموت بلا فائدة





Membrane potential is closer to the resting potential with increasing distance from the depolarization site.



1. صذا الخط المتقطع يمثد حاد ال membrane potential قبد قدوم ال stimulus حيث انه كان stimulus والي هي هون resting state and الدكتور قال ما رح اشدحه بس هذا مقارنة بين خصائص ال action potential و خصائص ال graded potential، بس شدح كم شفلة على سبيل المثال و هسا رح اكتبهم و الي مش واضح مني رح اوضحه

و هذا شيء اكيد انه رح يكون بس على أطراف العصبونات (الذوائد) ، لأنه هو بس رح ينقل action potential عشان يفعل ال synapses الإشارة العصبية من عصبون إلى آخد في ال postsynaptic عشان يفعل الكلام عادي اصبر لحد ما تخلص سلايد في العصبون الآخر postsynaptic (اذا مش فاهم هذا الكلام عادي اصبر لحد ما تخلص سلايد ٢٣)

Stimulated by neurotransmitters

Comparison of graded potentials and action potential

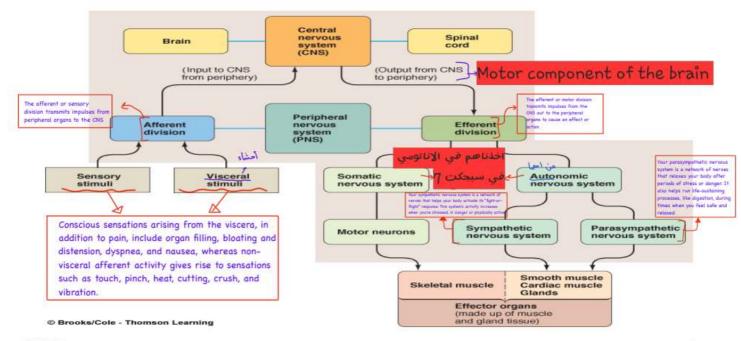
Characteristic	Graded Potentials	Action Potentials		
Origin	Arise mainly in dendrites and cell body (some arise in axons)	Arise at trigger zones and propagate along the axon.		
Types of channels	Ligand-gated or mechanically gated ion channels.	Voltage-gated channels for Na+ and K+.		
Conduction	Not propagated; localized and thus permit communication over a few micrometers.	Propagate and thus permit communication over long distance.		
Amplitude	Depending on strength of stimulus, varies from less than I mV to more than 50 mV.	All-or-none; typically about 100 mV. graded المسبة عني سريع بالنسبة لل Shorter, ranging from 0.5 to 2 msec.		
Duration	Typical longer, ranging from several msec to several min.			
Polarity	May be hyperpolarizing (inhibitory to generation of an action potential) or depolarizing (excitatory to generation of an action potential).	Always consist of depolarizing phase followed by repolarizing phase and return to resting membrane potential.		
Refractory period	Not present, thus spatial and temporal summation can occur.	Present, thus summation cannot occur.		

هذه قنوات ايونات لكنها ليست حساسة لفرق جهد مثل الي درسناها ، بل حساسة لاشياء ميكانيكية ، مثل ال cochlea في اذنك (القوقعة) هذه بسبب موجات الصوت تفتح قنوات الايونات فيها و ليس بسبب الايونات فيها و ليس بسبب الايونات فيها و ليس بسبب

و هذا سبب انه ال graded بقدرش يكمد potential بقدرش يكمد طريقه و ينتشر ذي ال action potential لانه لا يعتمد على قنوات ايونات حساسة لفرق الجهد الكهربائي تنتشر على طول محود العصبون

Functional organization of the nervous system

فيه كلام



3/27/2023

Functions of the Nervous System

هذا تلخيص لسلايد ع (

Sensory

Sense changes through sensory receptors

Motor

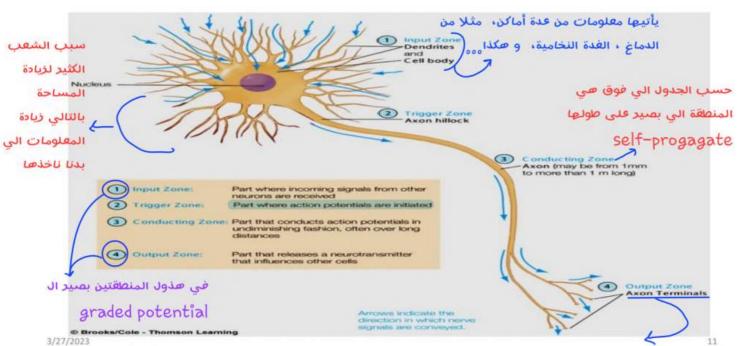
Respond to stimuli

Integrative

- Analyze incoming sensory information, store some aspects, and make decisions regarding appropriate behaviors
- The nervous system (1) receives millions of bits of information from the different sensory nerves and sensory organs and then (2) integrates all these to (3) determine responses to be made by the body.

ذي بمعنى يترجع هذه الإشارة التي جاءت و يفهم شو تعنى ليقوم بالإجداء المناسب

Functional component of a neuron



يوجد فيهم ال neurotransmitters، فلما يوصل ال action potential إلى ال neurotransmitters في action potential حج يتلقى ال graded هذه الإشارة و يقوم بتوليد terminals في العصبون التالي عن طريق تحريد النواقل العصبية

11/47

وحدة الوظيفة الإساسيةفي ال central nerve system neuron هي الخلية العصبية ، في اصغر مستوى تنظيمي وظيفى

General organization of the nervous system Functional classifications of neuron

The basic functional unit of the nervous system is the NEURON. The CNS contains more than 100 billion neurons. Neurons mainly function to store, communicate, and integrate informations

Sensory/afferent neurons

Inter/association neurons)

Motor/efferent neurons

Central nervous system Peripheral nervous system صدا مثال على ال peripheral Cell body Cell body Receptor Axon (central process) (peripheral process) Interneurons ينقل الإشارة للدماغ او لفيره Afferent neuron يتلقى إشارة من الدماغ او فيره لينقلها Efferent neuron إلى المكان المطلوب للتنفيذ Axon Axon terminal

The ones which carry information from the peripherals to the nervous system ولا الأعماب الي لما تلمس بخلوك تحس الأشياء

classification of nuerons

هاي بتكون موصولة بالعصلات و لما الدماغ يعطي امد بالحدكة هو حرح ينقل اله action potential للعصلة عشان تنقبض

هذول الأعصاب الي بكونوا بنص الطريق ما بين الأعصاب الطرفية و الحركية إلى الدماغ ،حتى ينقلوا الإشارة تاعت الدماغ للعصب الحركي او إشارة العصب الطرفى للدماغ

اليد للدماغ ، بل هي تشابكات

فصبية

Synapses and Signal Transmission

- A synapse is the junction between neurons or between a neuron and an effector like skeletal muscles
- Neuronal synapse is a junction between an axon terminal of one neuron, known as the presynaptic neuron, and the dendrites or cell body of a second neuron, known as the postsynaptic neuron. This junction allows the transmission of nerve action potential (or nerve impulse) from one neuron to the next.
- Types of synapse).

بشكل عام همه نوعين و

Electrical Synapse

electrical 91 chemical

Gap junctions connect cells and allow the transfer of information to synchronize the activity of a group of cells

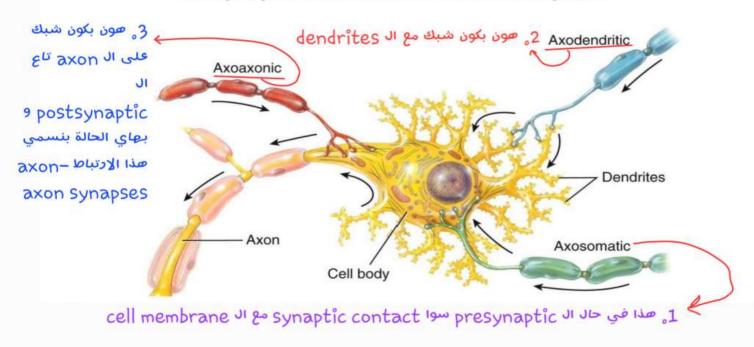
- Chemical Synapse
- One-way transfer of information from a presynaptic neuron to a postsynaptic neuron
- The primary type of synapses in the nervous system

هس الواحد اذا اجاه pain و بده يحس فيه ، يمشي التياد بواسطة (Synapses) من مكان الإصابة إلى الدماغ ليتم ترجمة هذه الإشارة على أنها ألم، لكن مثلا في حال بدي امشي رح تيجي ال signa من الدماغ إلى ليتم ترجمة هذه الإشارة على أنها ألم، لكن مثلا في حال بدي امشي رح تيجي ال Spinal cord لكي تقوم العصلة بالحركة

Types of Synapses Electrical synapses Functional anatomy اغلب الجسم من هذا معلومة وفي الدماغ لا يوجد electrical ، يوجد عندنا chemical A Chemical synapse B Electrical synapse Synaptic cleft synaptic delay 00 gap (20-40 Å) Synaptic cleft (200-300 Å) Functional contact Physical contact Ionotrop recepto Cellular response Membrane potential Biochemical cascades · Regulation of gene expression

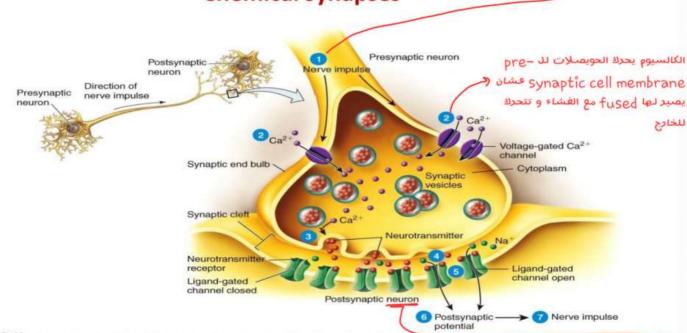
2. The nature of the channels

Structure of a chemical synapse and types different patterns of synaptic connection in nervous system



voltage gated Ca2+ channels العمد activation بيعمد action potential ال

Steps of synaptic transmission and Signal Transmission at a Chemical synapses

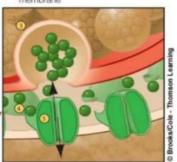


ح ارتباط النواقد العصبية في المستقبلات رح يفتح القنوات و يدخد الصوديوم (يزيد نفاذية الصوديوم) و دخود الصوديوم سوف يسبب depolarization ، و اذا كان ال depolarization كاف (دخود الصوديوم سوف يسبب postsynaptic cells ، و اذا كان ال sufficient) عشان يخلي اله postsynaptic cells يدتفع جهدها لجهد العتبة رح يصيد أnitiation an action potential

Steps In Synaptic Transmission



- An action potential is propagated to the terminal of a presynaptic neuron
- Ca²⁺ enters the synaptic knob
- Neurotransmitter is released by exocytosis into the synaptic cleft
- Neurotransmitter binds to receptor sites on the postsynaptic neuron
- Specific ion channels open in the subsynaptic membrane



النواقل العصبية لما ترتبط بالمستقبلات ما بصير تصل مرتبطة، بالتالي رح يتم إذالتها في ٣ طرق

Termination of action of neurotransmitters (Removal of Neurotransmitter)

1. Diffusion

2. Enzymatic degradation

Reuptake into presynaptic neurons

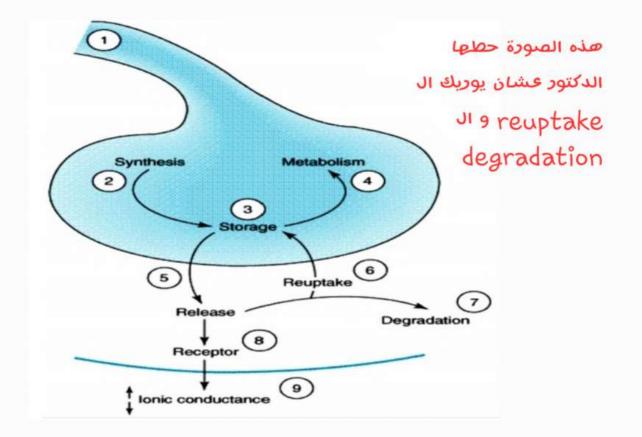
او سب ، presynaptic سب ، presynaptic

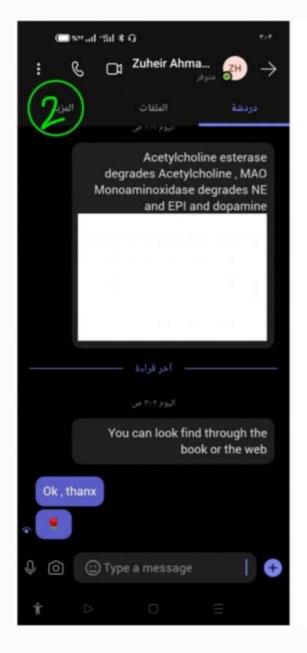
التركيز لإنه هذا انتشار

ممكن تيجي انزيمات معينة

تحطم النواقل العصبية

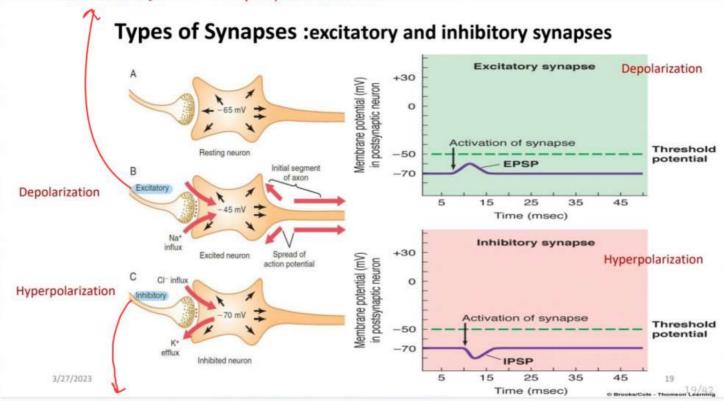
الي هي انه ترجع هذه النواقد لمكانها عن طريق active transport mechanism و هذه آلا reuptake السمها mechanism







اذا كان العصبون الي حامل الإشارة (presynaptic) إلى ال graded يقوم بإحداث عملية التحفيذ (يعني ال synaptic potential الي طالع منه لل postsynaptic يقوم بعملية excitatory فإننا نسميه (depolarization ا



اذا كان العصبون الي حامل الإشارة presynaptic إلى ال graded يقوم بإحداث عملية التثبيط ، (يعني ال synaptic potential الي طالع منه لك postsynaptic يقوم بعملية ال inhibitory ميسن نسميه (hyperpolarization

Excitatory Postsynaptic Potential (EPSP)

This potential is generated when transmitter acts on the membrane excitatory receptor to increase the membrane's permeability to Na^{\star} . The rapid influx of positively charged Na+ to the interior neutralizes part of the negativity of the resting membrane potential. In the figure the resting membrane potential has B increased in the positive direction from -65 to -45 mV, to a less negative value. This less negative value (closure to the threshold) is called the excitatory postsynaptic potential (or EPSP). The mathematical value is EPSP= New membrane C potential-RMP (resting membrane potential). The Cl- ir value is always positive (i.e. millivolts more positive than the resting value). It is +20 mV in this example. 3/27/2023

لما تطرح سالب كبير من سالب صفير بطلع الجواب موجد

-45-(-60)=-45+60=15

أين يتم توليد ال action potential في ال postsynaptic؟ في ال axon hillock و هي نفسها بنقدر نسميها :

Initial segment of the axon

Where the axon leave the cell body (soma)

واضح الكلام ، بحكيلك فادة انه لا يكفي فصبون واحد presynaptic فشان يدفع جهد الداحة تاع ال postsynaptic إلى العتبة ، لاذم اكثد من فصبون يأثدوا في نفس الوقت و نعمد لد graded تافهم فملية Summation فشان يصيد فندي تأثيد

Excitatory Postsynaptic Potential - EPSP (cont.):

- Discharge of a single presynaptic terminal can only generate a small EPSP. In the spinal cord, discharge of as many as 40 to 80 terminals may be needed to bring large EPSP all the way up to -45 millivolts. This simultaneous discharge occurs by a process called summation.
- When the EPSP rises high enough in the positive direction, an action potential initiates in the initial segment of the axon where the axon leaves the neuronal soma (i.e. axon hillock or the trigger zone).
- ☐ The membrane of the hillock segment has seven times as great a concentration of voltage-gated sodium channels as does the soma and the dendrites and, therefore, can generate an action potential with much greater ease than can the soma.
- An EPSP between +10 and +20 mV will elicit an action potential in the axon initial segment.

الجهد الي جاي من presynaptic و الي دح لي الي دح لي presynaptic ينشئ لي EPSP لاذم يدفع جهد الداحة بمقداد +10 إلى +20 عشان يتكون عندي action potential

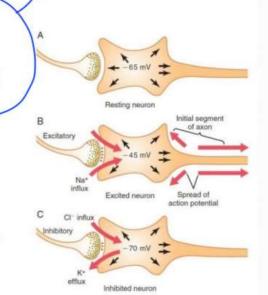
هون بحكي لك انه ال _{axon} الي بصير عليه ال action potential بكون كمية قنوات الصوديوم فيه تساوي سبعة أضعاف القنوات في ال soma و ال dendrites لذلك تلاقي انه الـ action potential فيه يتضاعف و يكبد و يصبح له تأثيد كافي لأداء الأمد المطلوب

Excited neuron

يفتح قنوات الكلود (سالب) عشان يعمد له ۱nflux (دخود للداخد) و يفتح قنوات البوتاسيوم (membrane potential (دخود للداخد) وبلخارج) بالتالي رح يصيد اله وfflux موجب) عشان يعمد مدائم (خدوج للخارج) بالتالي يمنع نشوء اله hyperpolarization سالب كثيد لدرجة اله hyperpolarization بالتالي يمنع نشوء

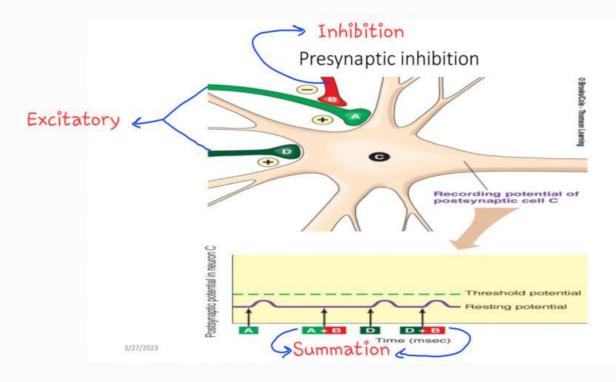
Inhibitory Postsynaptic Potential - IPSP:

- □ The inhibitory synapses open mainly chloride channels, allowing easy influx of chloride ions.
- Also opening potassium channels will allow positively charged K+ to move to the exterior and will also make the interior membrane potential more negative.
- □ Thus, both Cl⁻ influx and K⁺ efflux increase the degree of intracellular negativity (i.e. hyperpolarization) → inhibitory postsynaptic potential (IPSP).
- □ In the diagram the value of the IPSP is -5mV. This means 5 millivolts more negative than normal, which inhibits transmission of the nerve signal through the synapse.



الي فهمته من هذا الكلام هو انه لما اقول لك انه قيمة ال IPSP=-5 ، يعني هو نذل بمقدار 5 عن جهد الداحة ، يعني جهد الداحة =-70 ، لو صار -75 معناها ال Value لل IPSP=-5mv

هون بحكيلك انه في نوع من ال cell soma بكون مش متصد مباشرة مع ال cell soma الي بده يسوي لها أدام أنه في نوع من ال excitatory synapses مع مده معمد معمد النوع بكون عامل excitatory synapses مع مده معمد النوع بكون عامل excitatory synapses مع العصبون المداد نقد شابك مع اله مده والعصبون المداد نقد الهائة له ، بينما عامل axosomatic synapses مع A ، و هذا النوع من ال nhibitory synapses بكون أدم أنه المده ويعملوا على تثبيط ال neurotransmitters التي يحددها النوم من ال neurotransmitters التي يحددها العصبون المداد ا



بعمل مكتوب في نقطه(3)

يعني قبد ما ال neurotransmitters تاعون A يوصلوا إلى A و يفعلوا اله action potential، رح تكون ال معني قبد ما ال A و مدن المستقبلات سوف يقوم GABA neurotransmitters وصلت إلى مستقبلاتها في B ، و ادتباطها في هذه المستقبلات سوف يقوم

GABA neurotransmitters

موجودة في B

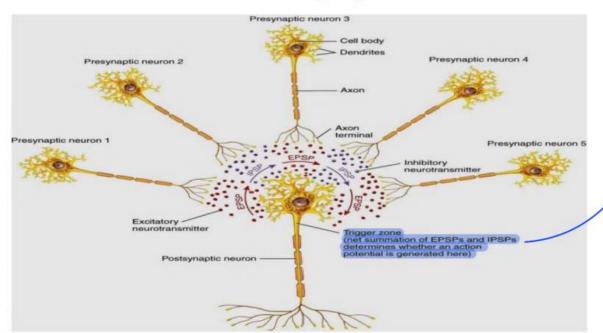
Presynaptic Inhibition

- It is an inhibitory input act on the presynaptic terminal before the signal ever reaches the synapse to make it less likely to communicate with postsynaptic neurons.
- Presynaptic inhibition is caused by release of an inhibitory substance, like GABA, acts on GABA receptors onto the outsides of the presynaptic nerve fibrils before their own endings terminate on the postsynaptic neuron. مستقبلات د GABA موجودة على GABA
- GABA receptors activation causes a chloride influx, which hyperpolarizes the cell that will cancel much of the excitatory effect of Na+ that also enter the terminal fibrils when an action potential arrives.
- Presynaptic inhibition occurs in many of the sensory <u>pathways</u> in the nervous system.

الآن وصول GABA لمستقبلاتها رح يجعد اله terminal fibrils تاعت B تفتح قنوات الكلور السالب و يدخل للازن وصول GABA لمستقبلاتها رح يجعد اله hyperpolarization و يلغي عمد اله اله اله اله اله اله تعلمنا مدانمة مدانمه عمد اله علما اله ع

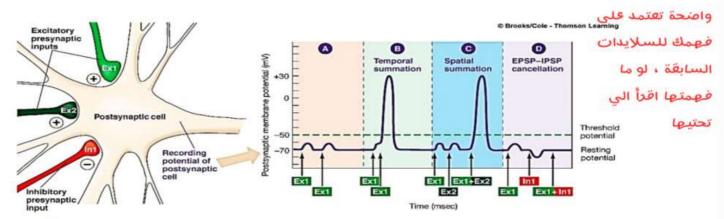
هون بحكيلك بالصورة هاي انهوفش خلية بتستقبد إشارة من عصبون واحد ، عادة بكون متصد فيها كثير presynaptic و لو كان عدد ال presynaptic الي عملوا excitatory synaptic معناها رح excitatory synaptic معناها رح يصيد لل postsynaptic عملية postsynaptic عملية ولوكانوا قد بعض ما رح يصيد شيء ،hyperpolarization

Summation of Postsynaptic Potentials



Summation of Postsynaptic Potentials

هذه الصورة جدا



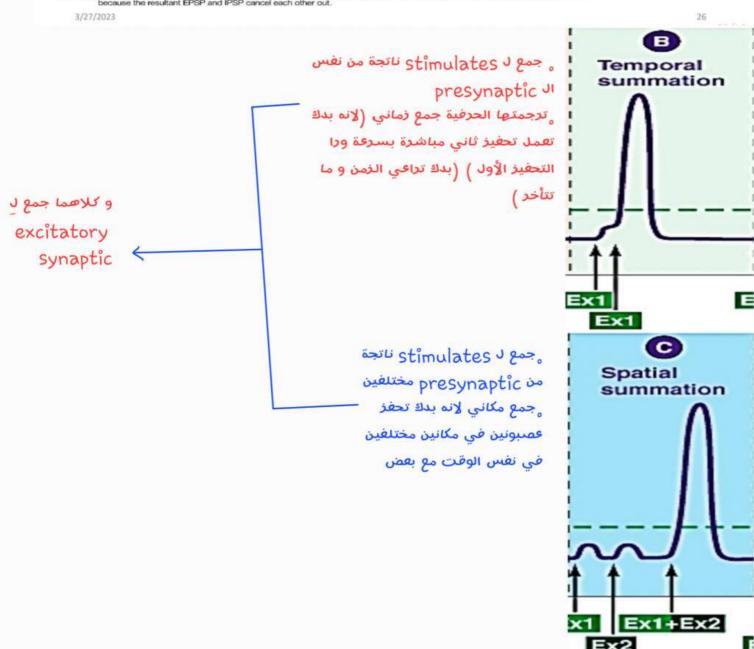
Panel (A) If an excitatory presynaptic input (Ex1) is stimulated a second time after the first EPSP in the postsynaptic cell has died off, a second EPSP of the same magnitude will occur.

Panel B If, however, Ex1 is stimulated a second time before the first EPSP has died off, the second EPSP will add onto, or sum with, the first EPSP, resulting in temporal summation, which may bring the postsynaptic cell to threshold.

Panel The postsynaptic cell may also be brought to threshold by spatial summation of EPSPs that are initiated by simultaneous activation of two (Ex1 and Ex2) or more excitatory presynaptic inputs.

Panel

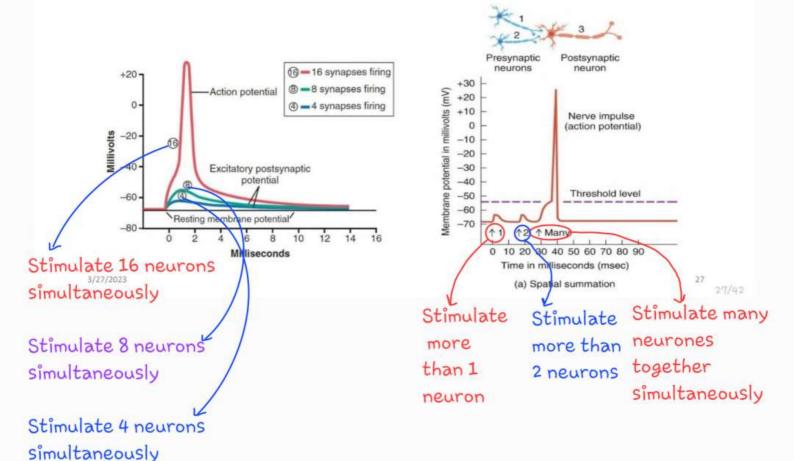
Simultaneous activation of an excitatory (Ex1) and inhibitory (In1) presynaptic input does not change the postsynaptic potential, because the resultant EPSP and IPSP cancel each other out.



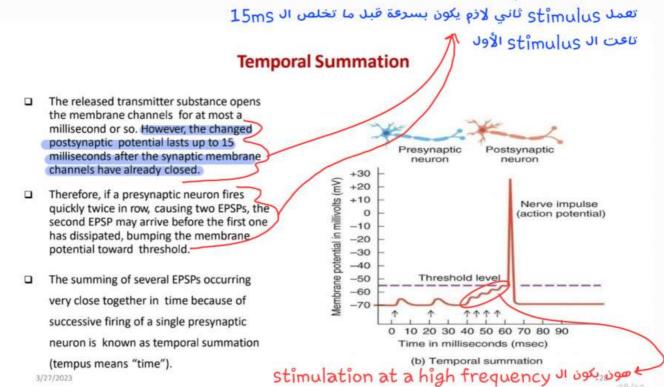
Spatial Summation

في نفس الوقت

is the summation of EPSPs originating simultaneously from several different presynaptic inputs (that is, from different points in "space").



هسا انت لما تسوي stimulus سواء خفيف او لأ ، هذا ال stimulus رح يعمل تفيد في ال stimulus عن stimulus عن طريق فتح القنوات) مدته 15ms ، بعد 15 خلص بتسكر، فإذا بدك تعمل stimulus ثاني لازم يكون بسدعة قبل ما تخلص ال stimulus



Neurotransmitters and chemical synapses

Neurotransmitters

- Chemicals synthesized, packed and released by presynaptic neurons
- · Diffuse across the synaptic cleft
- and interact with postsynaptic neurons membrane receptors
- Neurotransmitters either excite the neuron or inhibit or or modify its sensitivity of the postsynaptic neuron
- Inhibition or excitations is determined by the neurotransmitter, its receptors and permeability changes in postsynaptic cell induced by neurotransmitter receptor interaction

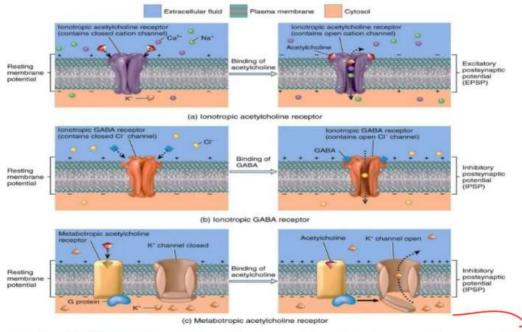
يسبيها

Structure of Neurotransmitter Receptors

- Neurotransmitters at chemical synapses cause either an excitatory or inhibitory graded potential
- Neurotransmitter/receptors/have two structures
 - Ionotropic receptors : gating ionic channels directly
 - Metabotropic receptors : activating a second messenger through G proteins coupled receptors

صداحة شرحهم واضح جدا ، اذا م*ا فهمت* شوف الصورة في السلايد الى ب*قد هذ*ا الدكتور ذكر معلومة و قال انه ال GABA لها كلا النوعين من ال receptors، واحد من نوع ionotropic كما في الصورة وواحد من نوع metabotropic من عندي لاحظ منوالرسمات انه ال acetylcholine كذلك

Ionotropic & Metabotropic Receptors



ارتباً neurotransmitters في ال receptor سيؤدي إلى إحداث تفاعلات كيميائية في ال neurotransmitters و تفير شكله فيفتح القناة المجاورة

الدكتور قال بديش تحفظوا هذه الصورة ، و لا كل ال receptors الي فيها ، بس القوا نظرة كيف عمل ال G-protein الدكتور قال بديش تحفظوا هذه الصورة ، و لا كل ال receptors الي فيها ، بس القوا نظرة كيف عمل ال Metabotropic Receptors

The second messenger system by which a neurotransmitter can affect the activity of postsynaptic cell

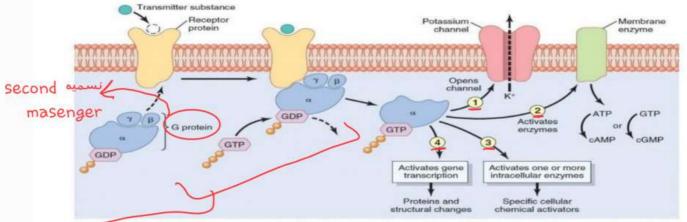


Figure The "second messenger" system by which a transmitter substance from an initial neuron can activate a second neuron by first causing a transformational change in the receptor that releases the activated alpha $\langle \alpha \rangle$ subunit of the G protein into the second neuron's cytoplasm. Four subsequent possible effects of the G protein are shown, including 1, opening an ion channel in the membrane of the second neuron; 2, activating an enzyme system in the neuron's membrane; 3, activating an intraceilular enzyme system; and/or 4, causing gene transcription in the second neuron. Return of the G protein to the inactive state occurs when guanosine triphosphate (GTP) bound to the α subunit is hydrolyzed to guanosine diphosphate (GDP) and the β and γ subunits are reattached to the α subunit.

قال الدكتور هون انه اله G-protein يقوم بعمل chemical reactions تعمل على تحويل ال GTP . و هذا اله GTP المرتبط ب G-protein سيقوم بعمل ٤ وظائف (من ١ ٢٥٠٠ ٤)

قال اكثر من هيك بديش تعرفوا

Types of CNS Receptors

Excitatory:

- · Ionotropic receptors:
 - · Nicotinic acetylcholine receptors
- · Metabotropic receptors:
 - · Muscarinic acetylcholine receptors
 - · Dopamine (D₁) receptors

· Inhibitory:

- · Ionotropic receptors:
 - GABA receptors
- Metabotropic receptors:
 - Opioid receptors
 - GABA_B receptors

هسا انتقال السيال العصبي يتأثر بدرجة الحموصة ، في حالة ال alkalosis (يعني يصير الوسط قاعدي كثير) دح تتحفز المعاب المعادة وومن الحالات التي بتسبب لي hyperventilation هي hyperventilation (الأعصاب بشكل كبير مما يسبب لها إجهاد، وومن الحالات التي بتسبب لي alkalosis هي CO2 هي المها تقلله الي هي فرط التنفس) و الي فيها دح تدخل كثير 02 و تخرج كثير CO2 لما تتنفس بسرعة ، و CO2 حمصني، لما تقلله المحديدة داخل جسمك و تتحفز أعصابك كثير

هذا عكس الي فوقها ، حالات مدضية بتعمل لي acidosis (تقلل درجة الحموضة pH) بالتالي دح تتثبط الأعصاب و ما تشتفل منيح كأنك متخدر

Notes on synaptic transmission

One way transmission

If an IPSP and an EPSP act on a neuron at the same time, these two effects can either completely or partially nullify each other. The summation of the membrane's EPSPs and IPSPs will determine the final membrane potential.

Synaptic potentials are graded and localized to site of stimulation

Exhibit summation

Repetitive neuronal stimulation at a rapid rate can develop **fatigue** of synaptic transmission. The mechanism of fatigue is mainly exhaustion or partial exhaustion of the stores of transmitter substance in the presynaptic terminals. The development of fatigue is a **protective mechanism** against excess neuronal activity.

- 1. Alkalosis greatly increases neuronal excitability (hyperventilation, which blows off CO₂ and elevates the pH, may precipitate an epileptic attack). بمعنى تقضى على ال Co² ، تقلل كميته كثير
- 2. Acidosis greatly depresses neuronal activity. In very severe diabetic or uremic acidosis coma develops
- Caffeine, theophylline (found in coffee, tea, and cocoa) increase neuronal excitability, presumably by reducing the threshold for excitation of neurons.
- 5. Most **anesthetics** increase the neuronal membrane threshold for excitation and thereby decrease synaptic transmission at many points in the nervous system.

3/27/2023

33/45

معدوف انه القهوة فيها كافيين بتخليك تصحيح لأنها تحفز الأعصاب زيادة عن الحد الطبيعي و، reducing the threshold for excitation of neurons و السبب انه بيعمد

هذه امثله على خصائص ووظائف ال _{Synapses}، قال لا تدققوا كثير عليها و أعطى بعض الشروحات لبعض الأمثلة، رح اكتبهم تحت في هذا السلايد More nots on synapses

Facilitatory and inhibitory signals from other areas in the nervous system can control the synaptic transmission at one site

The synapses perform a selective action, often blocking weak signals while allowing strong signals to pass.³

- The synaptic function include:
 - Blocking the transmission of impulses from one neuron to the next.
 - Amplification of signal: Changing the impulse from a single into repetitive impulses.
- (3) May integrate the impulse with impulses from other neurons to cause highly complicated or sophisticated patterns of impulses in successive neurons.
- Synapses have been suggested to play a role in memory and learning, behavior etc.

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1) If I want to flex my arm, the excitatory inputs will go to the flexor muscle, and the inhibitory inputs will be filter (block) whereas the excitatory inputs that go to triceps will block and the inhibitory inputs will go, and by this synaptic selective system you can flex your arm

مرات و هي الإشارة نازلة من النماغ و بده يصنخها عشان يصيد لها تأثيد قوي رح يمدرها في عصبون في نهايته terminals الكثيرة رح تتفدع فيها الإشارة إلى في نهايته terminals الكثيرة رح تتفدع فيها الإشارة إلى كثيد عصبونات ، بالتالي و

One presynaptic neuron will stimulate many postsynaptic neurons one presynaptic neurons 4 الدكتور عنهم اي مثال و لا قرأهم حتى بس همى واضحات خاصة له بتحكي انه مافك عبارة عن تجمع اعصاب مع بعض تقوم بعمليات الحفظ و التعلم و هكذا

شدح مني انا لنقطة 3 و انه من خصائص ال Synapses انه يجمع لك ال mpulses أمنوامثد من neurons و يصبها في neuron واحد (عكس نقطة 2 ، يعني عدة تفرعات تؤدي إلى عصبون واحد) و هذا يخلق لك أنماط متعقدة من ال impulses (لأنها مجموعة impulses مع بعض اكيد دح تكون قوية و معقدة لوظائف كبيرة و معقدة انا هيك فهمت هاى النقطة

من هون إلى سلايد ال neuropeptides كله انثلة على النواقد و أسمائها و كيفية عملها ، يعني كله حفظ للأسف ، الدكتور قال بديش تحفظوا كل الإسماء ، و قعد يقرأ قراءة عامة في الأسماء، و ما حدد مين The Neurotransmitters

- More than 50 chemical substances function as synaptic transmitters.
- These chemical substances can be grouped into; Small- molecule (rapidly acting transmitters) and Neuropeptide (slowly acting transmitters or growth factors).
- The small-molecule, rapidly acting transmitters cause most acute responses
 of the nervous system, such as transmission of sensory signals to the brain
 and of motor signals back to the muscles.
- The neuropeptides usually cause more prolonged actions, such as long-term changes in numbers of neuronal receptors, long-term opening or closure of certain ion channels, and possibly even long-term changes in numbers of synapses or sizes of synapses (Synaptic plasticity).

Small-Molecule, Rapidly Acting Transmitters:

- These types of transmitters are synthesized in the cytosol of the presynaptic terminal and are
 absorbed by means of active transport into the many transmitter vesicles in the terminal.
- Small molecule transmitters are **continually recycled** and used over and over again. و يتم presynaptic
- Acetylcholine, Norepinephrine, and Nitric oxide are examples of such transmitters.
- Acetylcholine (Ach): in most instances, has an excitatory effect; however, it is known to
- have inhibitory effects at some peripheral parasympathetic nerve endings (as in the heart).
- Norepinephrine (NE) is synthesized within the vesicle from dopamine the enzyme dopamine β
 hydroxylase is present in the vesicle. NE in many areas within the CNS, activates excitatory receptors,
 but in a few areas, it activates inhibitory receptors instead.

Small-Molecule, Rapidly Acting Transmitters

- Dopamine, Glycine, and GABA (gamma-aminobutyric acid) are inhibitory transmitters. <u>GABA is the primary inhibitory</u> neurotransmitter of the CNS. It functions as a CNS depressant.
- Glutamate is secreted by the presynaptic terminals in many of the sensory pathways. Glutamate is the principle excitatory neurotransmitter of the CNS.
- Serotonin (or 5-hydroxytryptamine) acts as an <u>inhibitor of pain</u> pathways in the spinal cord, and an inhibitor action in the higher regions of the nervous system. Serotonin is involved in <u>mood control</u>, <u>appetite control</u>, <u>and nausea</u>. Perhaps it even <u>causes sleep</u>.
- Nitric oxide is a gas and is not preformed and stored in vesicles in the
 presynaptic terminal as are other transmitters. Instead, it is synthesized almost
 instantly as needed and then diffuses out of the presynaptic terminals over a
 period of seconds rather than being released in vesicular packets. It diffuses
 into the nearby postsynaptic neurons and changes intracellular metabolic
 functions that modify postsynaptic neuronal excitability

The Neuropeptides Slowly acting transmitters

cell body.
 The Golgi apparatus packages the neuropeptide into minute transmitter vesicles that are released into the cytoplasm. Then the transmitter vesicles are transported all the way to the tips of the nerve fibers by axonal streaming of the axon cytoplasm (axoplasmic flow), traveling at the slow rate of only a few centimeters per day.

☐ They are synthesized as integral parts of large-protein molecules by ribosomes in the neuronal

- Much smaller quantities of neuropeptides than of the small- molecule transmitters are usually released at the neuronal terminals in response to action potentials.
- □ Neuropeptides are generally a <u>thousand or more times as potent as the small-molecule transmitters</u> and they often cause much more prolonged actions including calcium channels, prolonged changes in the metabolic machinery of cells, prolonged changes in activation or deactivation of specific genes in the cell nucleus, and/or prolonged alterations in numbers of excitatory or inhibitory receptors.
- ☐ Some of these effects last for days, but others last perhaps for months

acids

Small-Molecule, Rapidly Acting Transmitters Class I Acetylcholine Class II: The Amines Norepinephrine Epinephrine Dopamine Serotonin Histamine Class III: Amino Acids Gamma-aminobutyric acid Glycine Glutamate Aspartate Class IV Nitric oxide

Neuropeptide, Slowly Acting Transmitters or Growth Factors Hypothalamic-Releasing Hormones Thyrotropin-releasing hormone Luteinizing hormone-releasing hormone Somatostatin (growth hormone inhibitory factor) **Pituitary Peptides** Adrenocorticotropic hormone **B-Endorphin** a-Melanocyte-stimulating hormone Projectin Luteinizing hormone Thyrotropin Growth hormone Vasopressin Oxytocin Peptides that Act on Gut and Brain Leucine enkephalin Methionine enkephalin Substance P Gastrin Cholecystokinin Vasoactive intestinal polypeptide Nerve growth factor Brain-derived neurotropic factor Neurotensin Insulin Glucagon From Other Tissues Angiotensin II Bradykinin Carnosine Sleep peptides Calcitonin

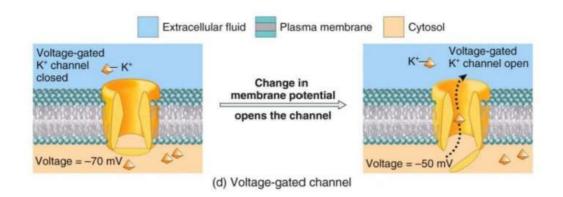
الدكتور جاب سيدة ؛ muscarinic receptors and nicotinic receptors، صداحة بعدفش وين مكتبوين و من وين جابهم ، هو جاب سيدتهم من دقيقة 41°55 إلى 43°15 و يمكن عدت هاي الجذئية ١٠ مدات ، لاقدر استوعب اسمهم شو هو ، و قال مهمات رح ييجي عليهم سؤال، و الي حكاه كان كالتالى ؟

Muscarinic receptors cns ، ، وجود في ، viscera organs , heart , stomach Nicotinic receptors

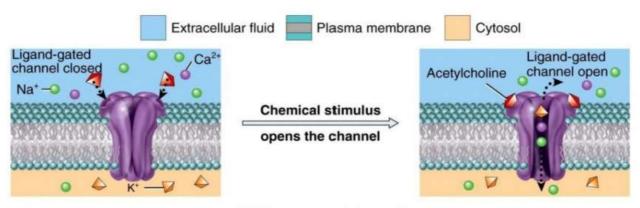
cns و neuro و neuscular junction و بتصيد و الله العضلية، الي بتطلع بين الخلايا العضلية، الي بتطلع من ال spinal cord و بتدوح إلى skeletal muscle الـ skeletal muscle

هاي الجزئية من الريكورد صداحة مش متأكد منها و قعدت ربع ساعة عليها ، و بحثت في قوقد مش ملاقي اشي ذي الكلام الي حكاه الدكتور و حتى كلامه عنهم كان شوي مخدبط ، فأرجوا انه ترجعوا تأكدوا من هاي الجزئية من الريكورد و اذا في فلط احكولي رجاءً

Voltage-gated channels respond to direct changes in membrane potential



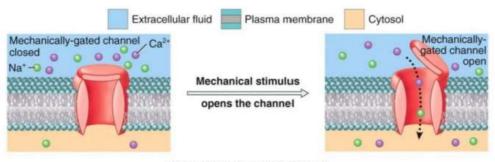
Ligand-gated channels respond to chemical stimuli (ligand binds to receptor)



(b) Ligand-gated channel

A graded potential occurs in response to the opening of a mechanically-gated or ligand-gated ion channel

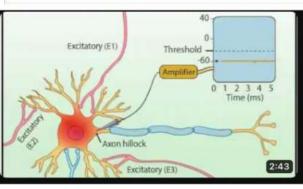
Mechanically-gated channels respond to mechanical vibration or pressure stimuli



(c) Mechanically-gated channel

A graded potential occurs in response to the opening of a mechanically-gated





EPSP & IPSP 115K views · 5 years ago

S Stens Biofeedback

Neurons communicate with each other mainly mainly through chemical messengers or neurotransmitters mainly neurons are generated. Neurotransmitter neuron, neurotransmitter. A pre-synaptic axon may form a synapse with either a dendrite, a cell body or the axon of a post-synaptic neuron giving rise to an axonal synapse, a synapse or an axon respectively. Chemical synapses exist not only between neurons but also between neurons. Nerve cells and target cell such as muscle or gland cellMore than a hundred neurotransmitters have been identified so far, most of which can be classified into classes according to their chemical structure. The main classes include: amino acids such as glycine, glutamate, aspartate, and GABA small peptides, called neuropeptides, such as beta-endorphin and substance P-monoamines, such as epinephrine, norepinephrine, dopamine, serotonin, and histamine.

Monoamino acids are essentially S amino acids with the acidic group removed and acetylcholine, the choline ester, is in its class alone

Neurotransmitters are synthesized in presynaptic neurons and stored in small sacs, called synaptic vesicles, at the axon terminal. Some of these vesicles are anchored to the plasma membrane, ready to release a neurotransmitter on demand. When an action potential reaches the nerve terminal, the resulting depolarization opens voltage-gated calcium channels, allowing an influx of calcium. Calcium causes vesicles to fuse with the plasma membrane, resulting in release of the neurotransmitter in a process known as exocytosis upon binding to its receptors on the postsynaptic cell., Some neurotransmitters open ligand-gated ion channels, causing direct changes in the membrane potential of the receiving neuron while others act through second reporter systems to exert their influence.

Some neurotransmitters are excitatory, others are inhibitory and for some the effect can be either excitatory or inhibitory depending on the receptor they bind to.

An example of an excitatory neurotransmitter is glutamate. Upon binding, glutamate receptors trigger the interconnected ion channels to open and allow positively charged ions to enter the cell, making them more positive, less polarizing, and therefore more likely to generate action potentials. On the other hand, GABA, a major inhibitory neurotransmitter, opens up ligand-gated chloride channels to allow entry of negatively charged chlorides, making the cell more negative and more polarized and thus less likely to generate action potentials.

Acetylcholine is a neurotransmitter that can be excitatory or inhibitory depending on the receptor present in the target cell at the neuromuscular junctions. Acetylcholine secreted by motor neurons binds to nicotinic receptors in skeletal muscle cells and stimulates them to contract. On the other hand, it inhibits cardiac muscle cells from by the M2 muscarinic receptor, which slows the heart rate, as part of the parasympathetic "rest-and-digest" response

The neurotransmitter binds to its receptor for only a millisecond or so, then passively diffuses from the synapse and is picked up by nearby astrocytes for recycling.

If the presynaptic neuron continues to release more neurotransmitters, the new molecules will bind and again activate the receiving neuron. If the presynaptic signal stops firing, the transmission will eventually stop. There are also mechanisms for removing neurotransmitters from the synaptic space to avoid overstimulation

Common mechanisms include: degradation of neurotransmitters by an enzyme present in the synapse and its re-uptake, whereby the neurotransmitter is transported back by a transporter protein to the presynaptic neuron for reuse



1. The human nervous system is capable of a wide range of functions. What is the basic unit of the nervous system?	
OA. Glial cell	
OB. Meninges	
OC. Neuron	
OD. Cerebrospinal fluid	

2. The neuron cell is made up of which of the following parts?
OA. Axon
○B. Dendrite
OC. Nucleus

OD. All of the Above

3. Neurons come in which different type(s)?
OA. Sensory
OB. Motor
○C. Skeletal
OD. A and B
4. How do neurons communicate with one another?
OA. Electrically
OB. Chemically
OC. Through weak, radio-wave-like impulses
OD. A and B
5) WHEN A NEUROTRANSMITTER BINDS TO A RECEPTOR LOCATED ON THE SURFACE OF THE PLASMA MEMBRANE, THE ACTION OF THE NEUROTRANSMITTER IS INHIBITED BY MANY WAYS SLICH AS

ENZYMATIC DEGRADATION WHICH INVOLVES:

- A) MAO
- **B) NORADRENALINE**
- C) ACETYLCHOLINESTERASE
- D) ACETYLCHOLINE
- E) A+C

Answers:

1) C

2) D

3) D

4) D

5) E



بالتوفيق

#النادي_الطبي

#معكم_خطوة_بخطوة



رسالة لكل حدا لسا زعلان على علامة الأورغانيك 🍅