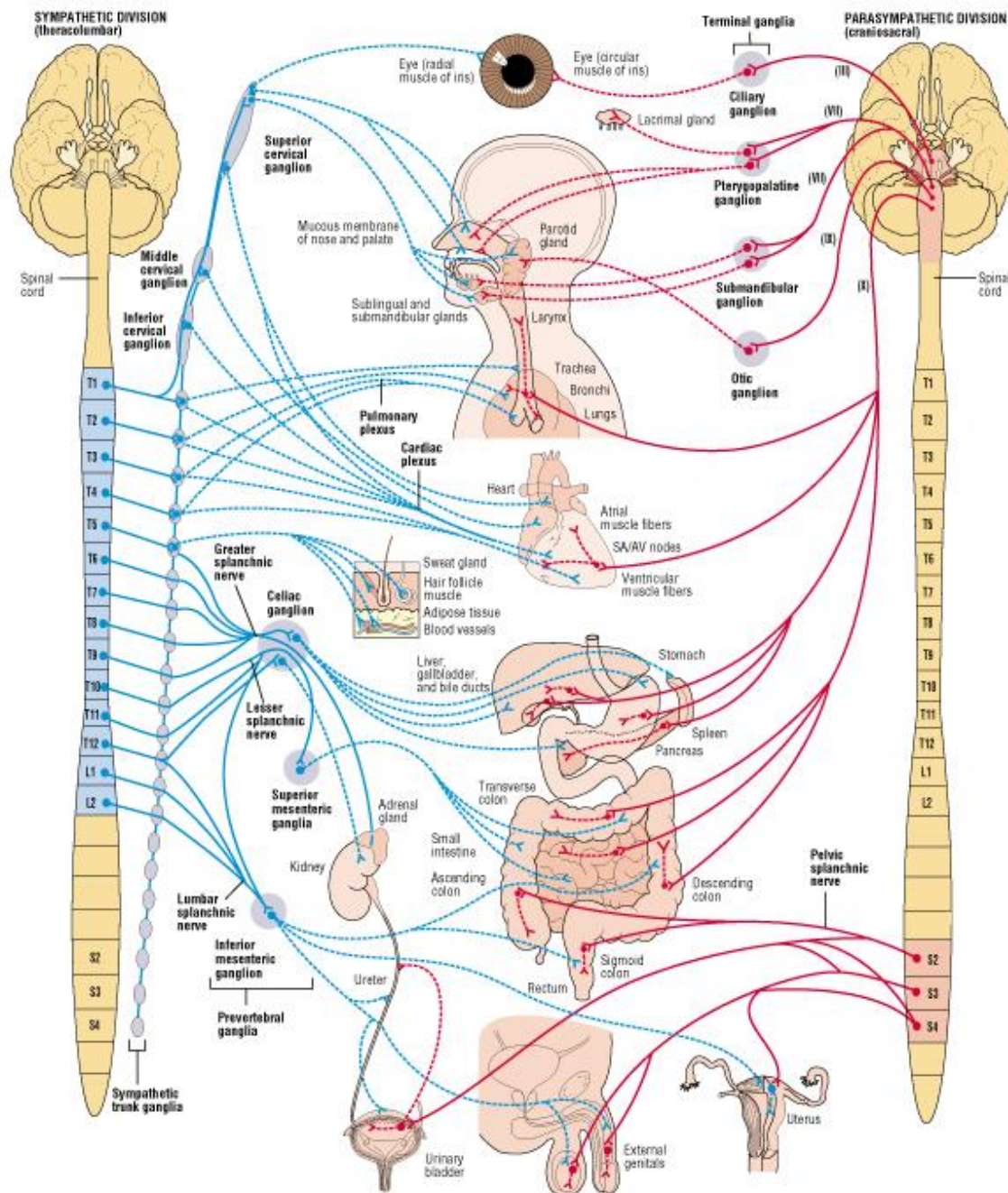


Autonomic Nervous System

Autonomic Nervous System

Ref: **Textbook of Medical Physiology**, Guyton, 14th Ed.:763-773, 13th ed.: 773-784. 12th ed: 729-738, and 11th ed. P748-760.

Fig.17.02



2 divisions
of
ANS

General functions

importance ↘

Control and Adaptation of body systems to
internal and/or external changes

Example of adaptation to external stimuli

Fight and Flight Reaction → Sympathetic.

in both situations

we need the activity of muscles.



All body systems are working to increase the delivery of O_2 towards these muscles.

① - Increase heart rate and force of contraction. → more delivery of O_2 → muscles.

- Widely dilated pupils. "mydriasis" (to see all details of that dangerous object).

- Pallor (pale of fear) as blood is directed to the skeletal muscle. (discoloration of skin)

- Goose pimples. = when the hair stands on the skin.

- Cold sweat. → at the base of hair follicles we have smooth muscles (pilomotor muscles) by activating them it cause the hair to stand up.

- Dry mouth.

many sweat glands are activated → (Cold) because of low distribution of blood towards the skin → so we are not heating that sweat.

Because we have decreased the activity of GIT, one of its glands is the salivary gland → decrease in salivation.

* The decrease of secretion happens Bcs of vasoconstriction → less fluids available for secretion

we have vascularization in skin → in this case we reduce / redistribute blood → more blood towards muscles

→ less blood for unnecessary tissues like skin.

↓
Vasoconstriction for vessels supplying skin

+ vasodilation for ones supplying muscles.

What division is involved in the control during fight or flight reaction???

The sympathetic division



ANS characteristics

①

Anatomical characteristics and
Synaptic^② organization of ANS

Fig.17.02

Origin :-

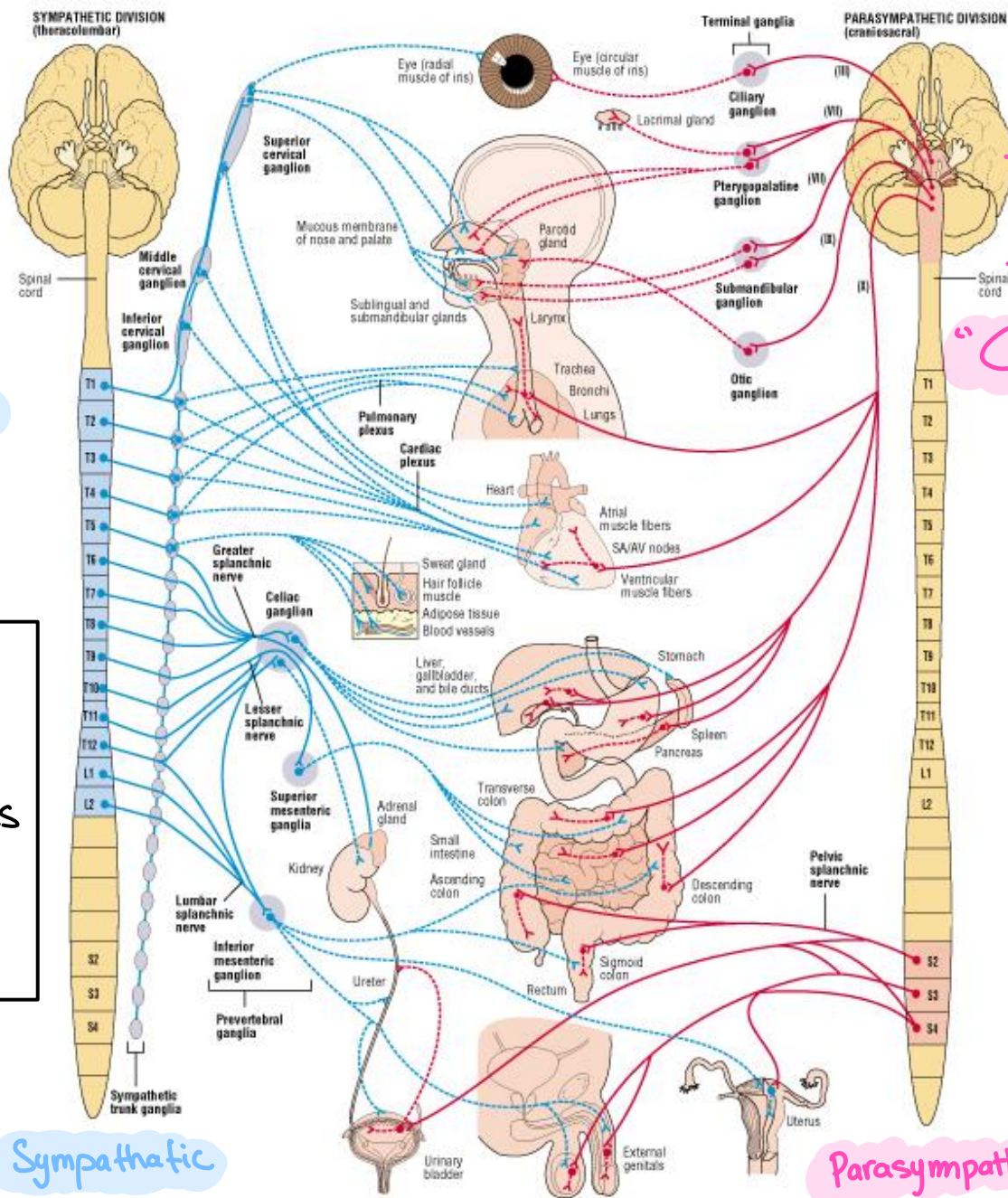
→ Spinal Cord

(T₁ → L₂)

"thoracolumbar"

Similarities :-

→ from the origin until the effector we have 2 neurons forming the arc of that division.



Origin :-

→ Base of the Brain

→ Sacral region.

"Craniosacral"

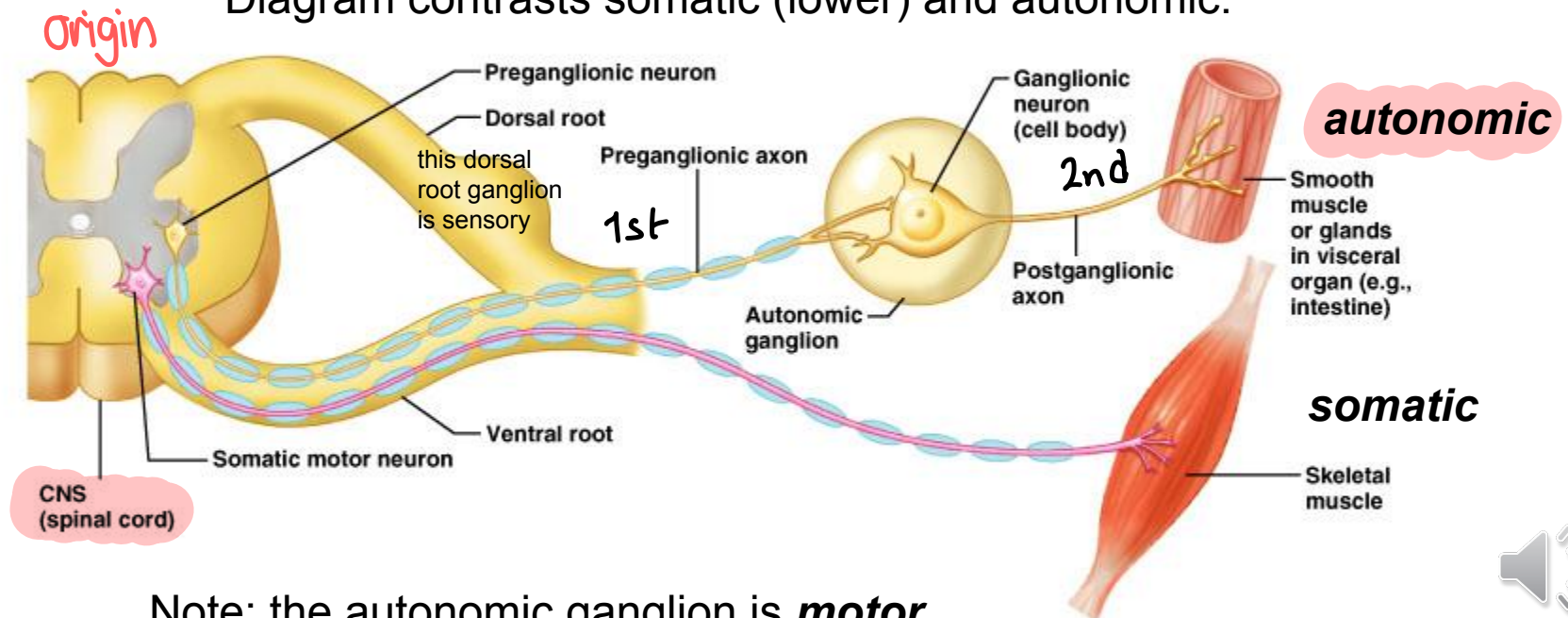
Sympathetic

Parasympathetic.

- Axon of 1st (*preganglionic*) neuron leaves CNS to synapse with the 2nd (*ganglionic*) neuron
- Axon of 2nd (*ganglionic*) neuron extends to the organ it serves

→ in somatic we have only one neuron forming the arc from the origin until the effector.

Diagram contrasts somatic (lower) and autonomic:

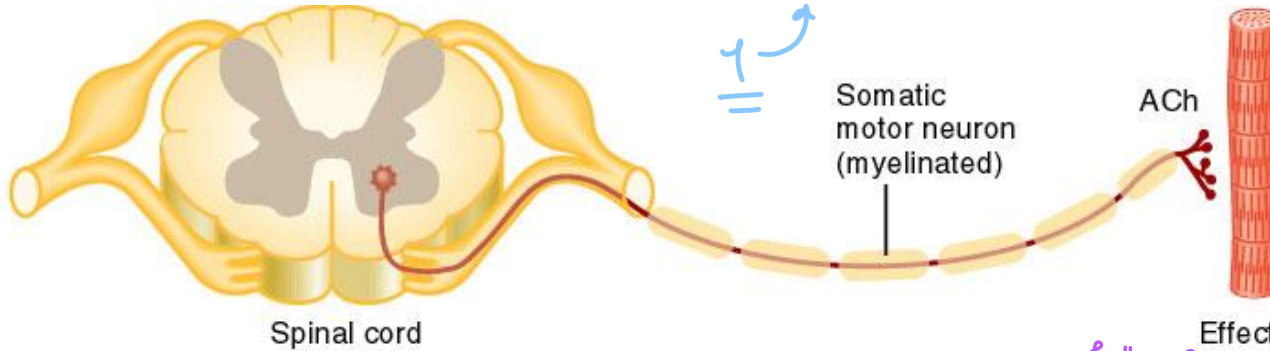


Note: the autonomic ganglion is *motor*



Fig.17.01

axon of 1st neuron reaches the effector organ



Somatic

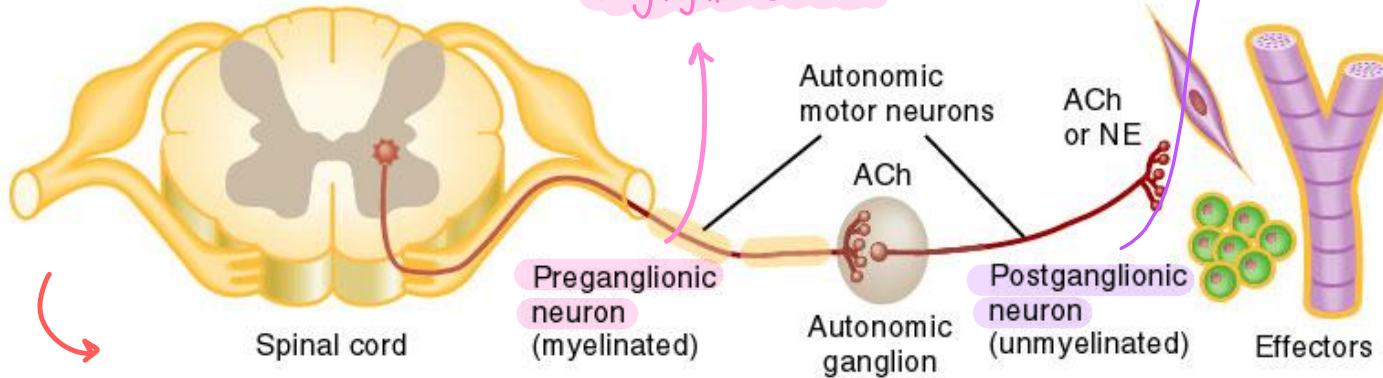
ACh: Skeletal muscle-contraction

2 nerves

→ axon of the 1st neuron is called Preganglionic fiber

axon of the 2nd neuron is called Postganglionic fiber

Autonomic



ACh or NE: Smooth muscle-contraction or relaxation
Cardiac muscle-increased or decreased rate and force of contraction
Glands-increased or decreased secretions

1st neuron synapsing with 2nd neuron at the level of a ganglion

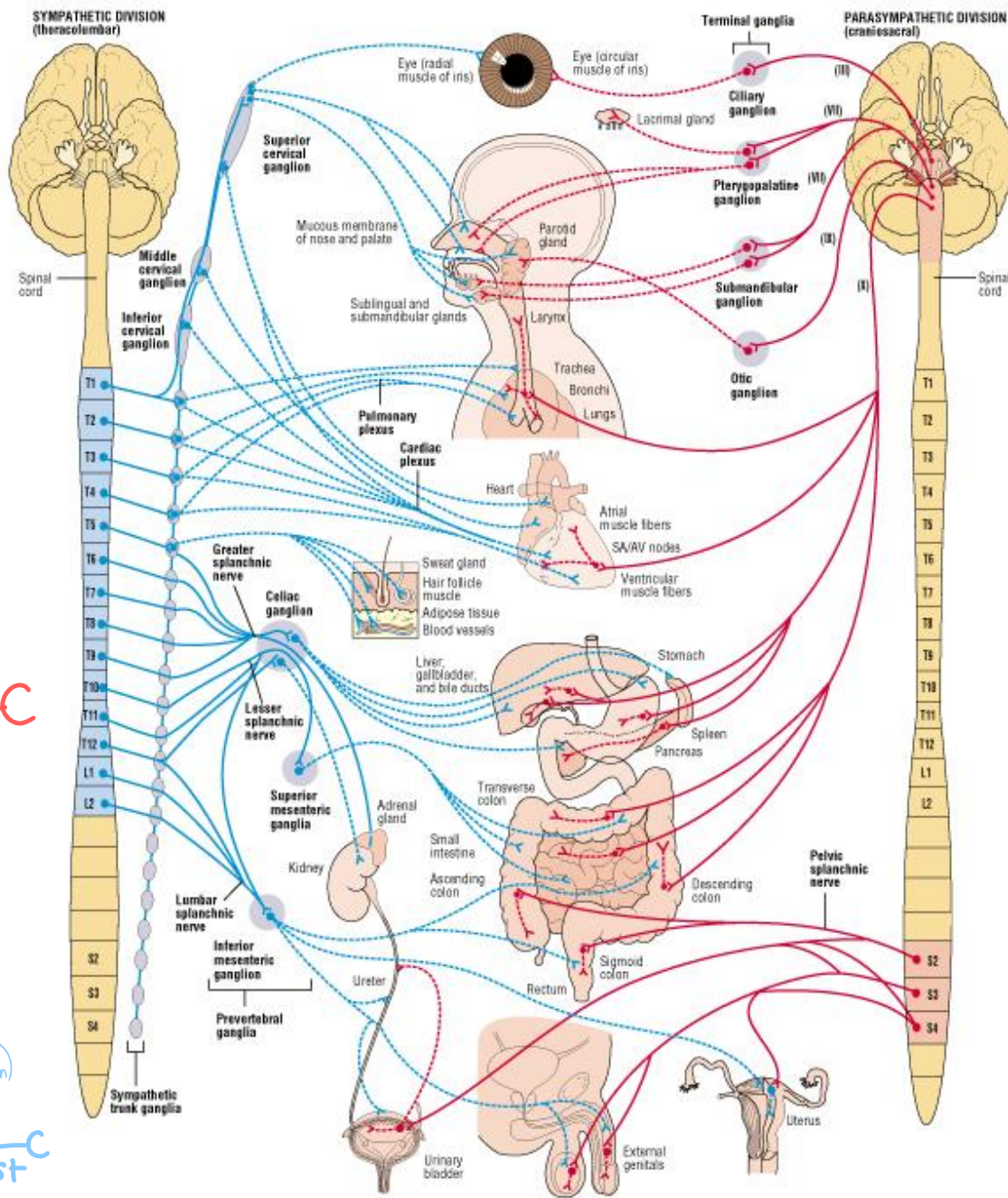
(b) Autonomic nervous system

17.01

↳ a ganglia : is a cluster of cell body of neurons which represents the 2nd neuron.



Fig.17.02



• differences :-

→ Sympathetic.

Pre ganglionic fibers

Short

Post ganglionic fibers

Long



→ Parasympathetic

Pre ganglionic fibers

Long

Post ganglionic fibers

Short (bated at the effector organ)



• Similarities :-

2 neurons forming the arc



Synaptic organization of ANS

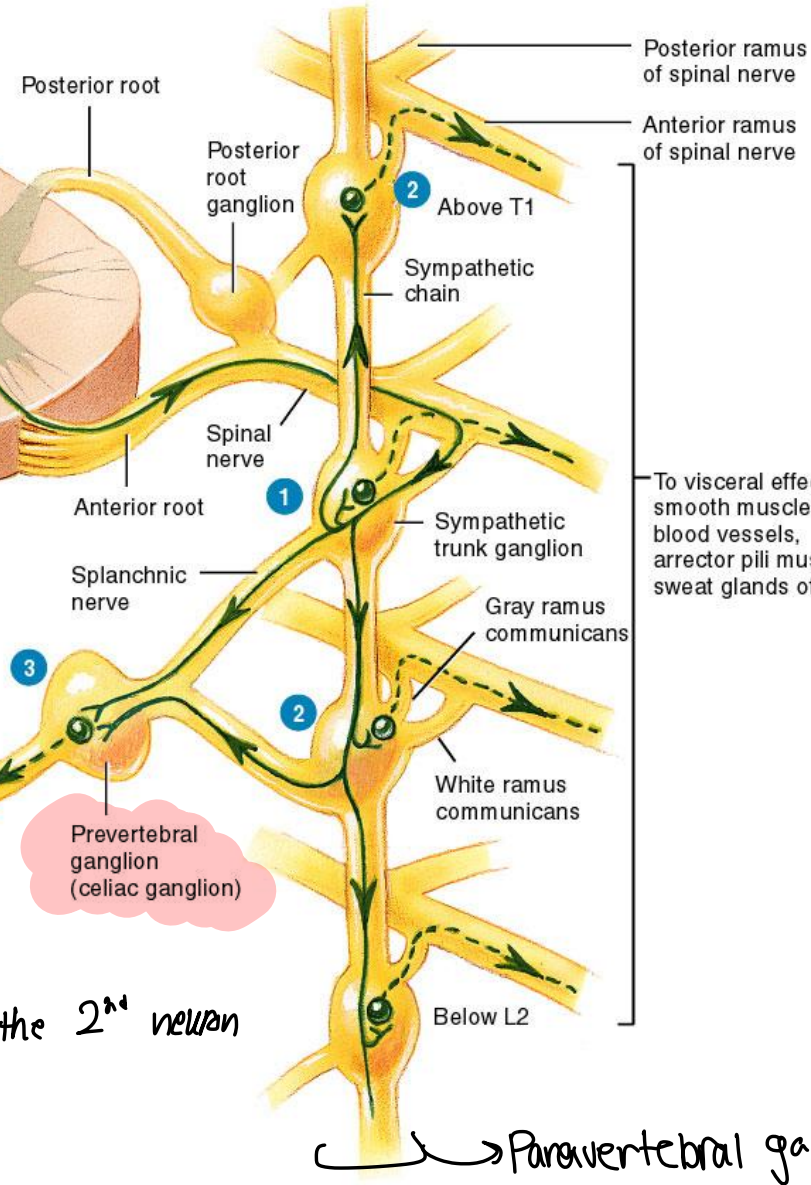
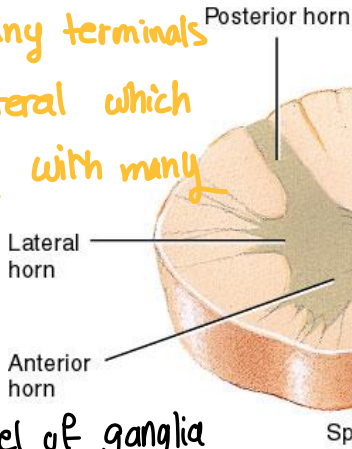
Convergence and Divergence in Sympathetic division



Divergence \rightarrow in Sympathetic

Fig. 17.05

\rightarrow pre ganglionic fibers are having many terminals and many collateral which are synapsing with many neurons



Posterior ramus of spinal nerve
Anterior ramus of spinal nerve
To visceral effectors: smooth muscle of blood vessels, arrector pili muscles, sweat glands of skin

\rightarrow at the level of ganglia at the segmental level

\rightarrow upper segmental level

\rightarrow lower segmental level.

Some also collaterals from Preganglionic fibers passing without synapse at this paravertebral ganglia \rightarrow to synapse with the 2nd neuron in the abdominal cavity.

\rightarrow the group of ganglia found inside abdominal cavity \Rightarrow Pre vertebral Ganglia.

— Preganglionic neuron
- - - Postganglionic neuron

Anterior view



Convergence

→ neuron located at the same segmental level → it can receive terminals or preganglionic fibers synapsing with that neuron coming from upper/lower/same segmental level of the sympathetic division to synapse with that neuron.

Collateral from many neurons synapsing with one neuron ⇒ Convergence.

* parasympathetic → we don't have that high divergence and convergence *

A difference between sympathetic + para in the synaptic organization.

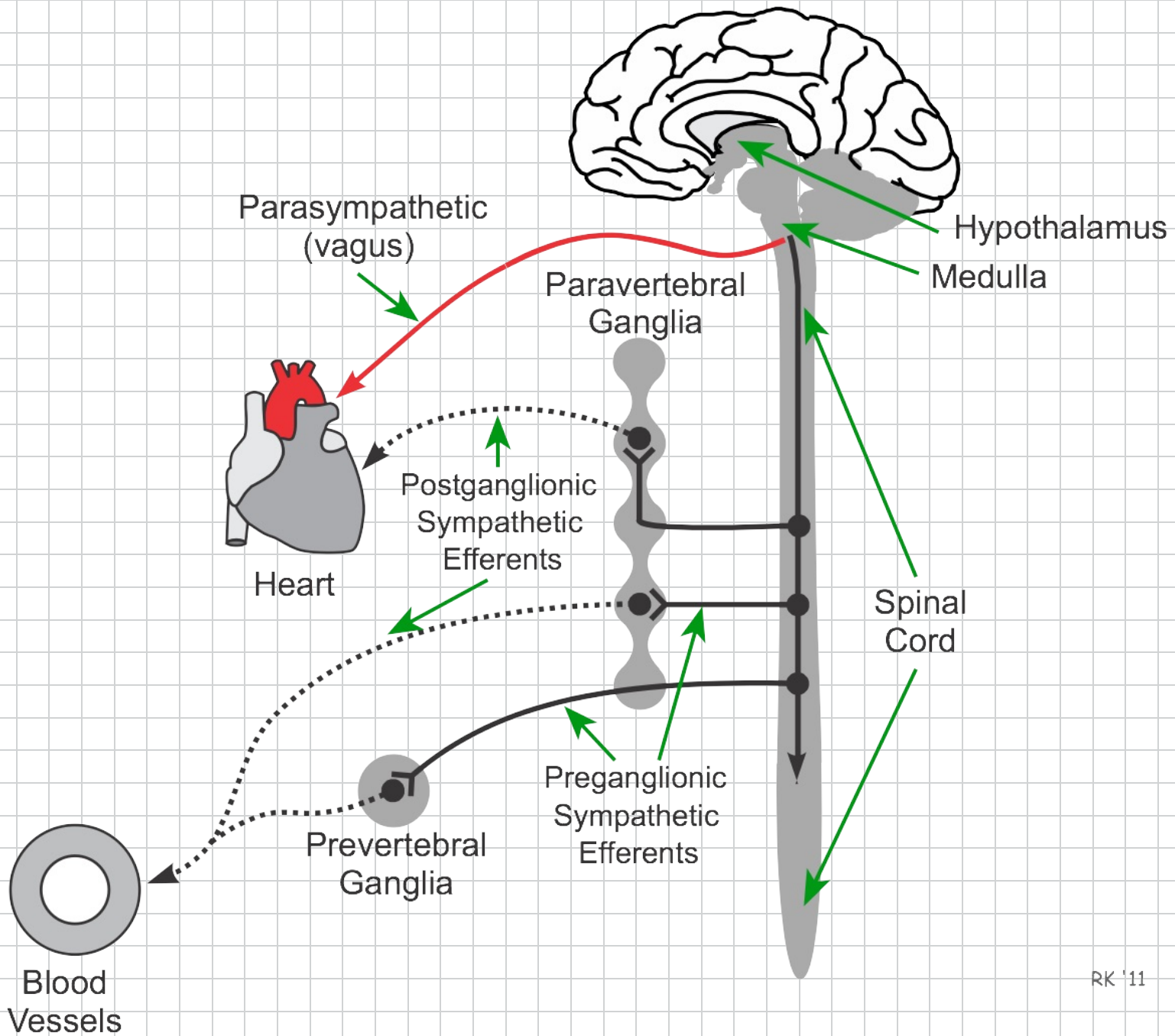
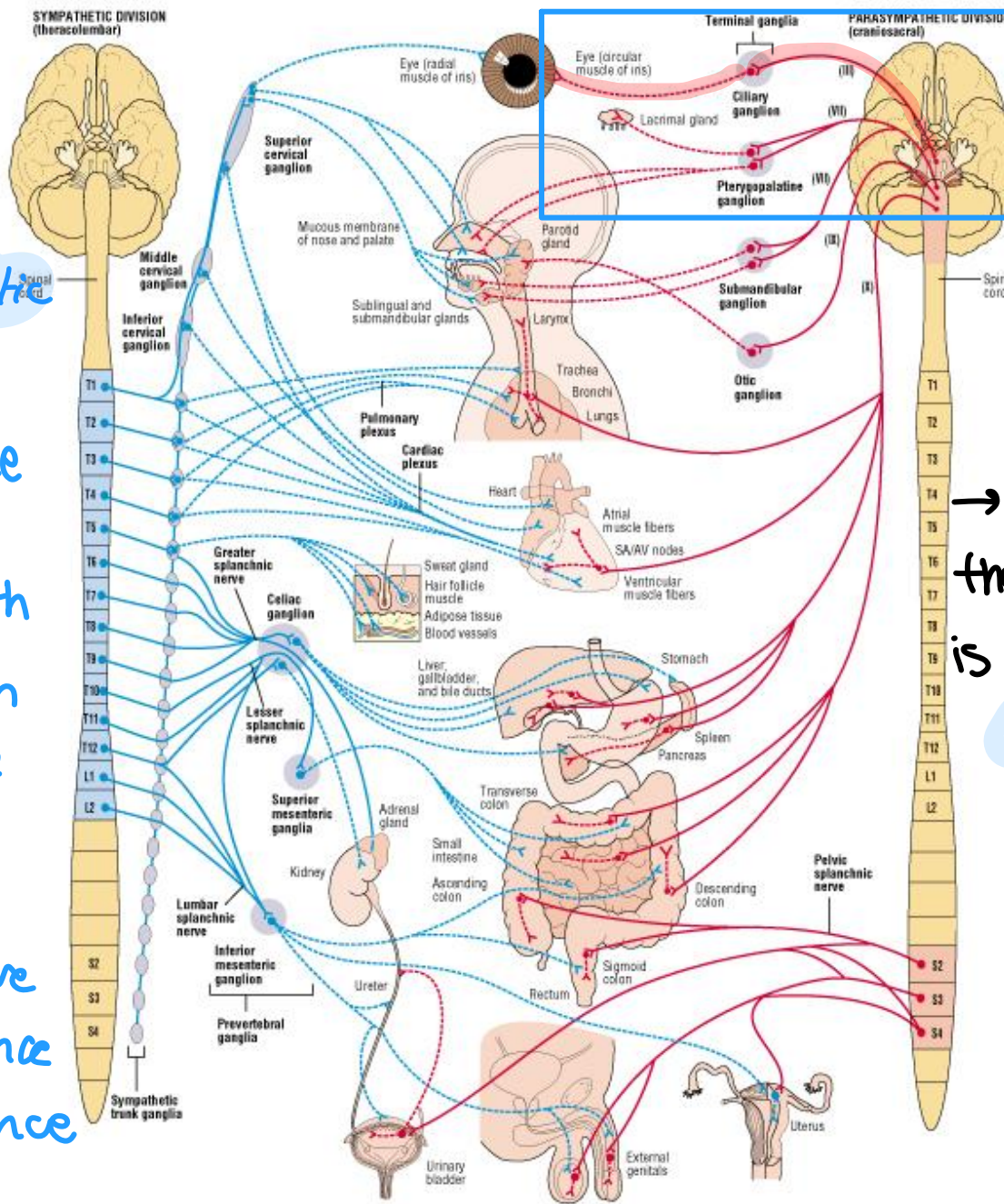


Fig.17.02



Divergence
+
Convergence
In Parasympathetic

↓
the axon of the
1st neuron
is synapsing with
the 2nd neuron
located at the
organ it self
↓
We don't have
that divergence
and convergence

neuron is
located at the
organ.

→ ratio between
the pre & post
is almost
1:1



Synaptic organization of ANS

What is the importance for having more Convergence and Divergence in the Sympathetic division?

- by stimulating the **sympathetic** system, because of that synaptic organization the reaction that you are getting is more **diffuse** reaction. [at fight or flight we are getting All the reactions. High sweating, increased Heart rate, dry mouth. ...]
- but in **Parasympathetic** → we have no convergence or divergence → the reaction for Para-sympathetic stimulation is more **localized**. [eating → activates stomach only, but no decrease in ♡ rate]

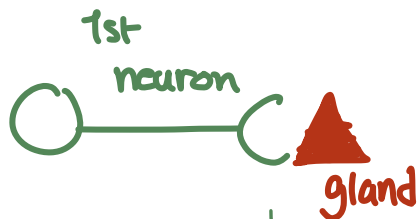


Adrenal gland is exception

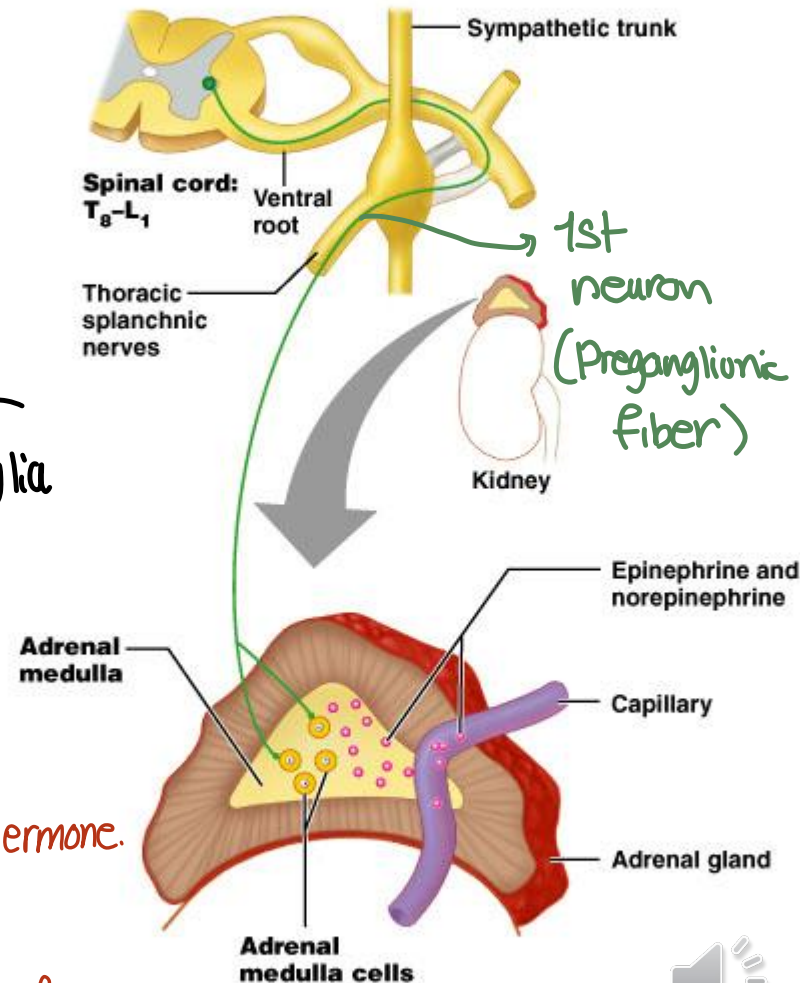
- Synapse in gland
- Can cause body-wide release of epinephrine

→ We can have some collaterals passing
Para vertebral ganglia → pre vertebral ganglia
→ suprarenal glands → Adrenal Medulla

↳ where we have endocrine cells, that releases
• epinephrine as a hormone.



↳ Synapsing with endocrine cells from suprarenal glands.



Physiological characteristics of ANS

- High speed of onset: there's no delay in getting response.
- Automatic nature: you can't control it, it is involuntary act
- Tonic activity: to get the control by sympathetic or parasympathetic stimulation.
↳ Certain level of activity ↓

for example :

→ neurons of the sympathetic divisions are generating 100 AP per min.

↳ tonic level of activity

we can get that tonic increased by generating more APs per time unit

OR decreasing the tonic activity by generating less APs per time unit.



Effects of **sympathetic** stimulation

- **Blood pressure** (blood vessels supplying skeletal muscle are major players). In addition to that the effect on heart also contributes in regulation of blood pressure.

- **Body temperature** by the sympathetic effects on cutaneous blood vessels and sweat glands.

↓
Vasodilation

↓
increasing Activity.

• vessels + sweat glands → widely distributed tissues in our body → for the Activity of widely distributed tissues → we need a division that can have **diffuse** effect = **Sympathetic**

→ Widely distributed tissues are Innervated only by sympathetic NS [Ex. Blood vessels / Glands]



Effects of **sympathetic** stimulation

- **Cardiovascular system**: effects on vessels will result in redistribution of blood by enhancing blood flow to skeletal muscle and reducing blood flow to skin and mesentery.
- **Effects on heart**: increasing cardiac output (volume of blood pumped per minute). → to deliver more oxygenated blood to muscles.
- **Respiratory system**: causes relaxation of bronchial muscle which result in bronchodilation. to get more Air flow towards the lung + more
- **Digestive system**: inhibition of motility and secretion. exchange of gases by the lung.
- **Metabolic effects**:
 - * Mobilization of glucose.
 - * Increased lipolysis.
 - * Increased metabolic rate. → to mobilize nutrients for the activity of muscles to provide these muscles during fight - flight by these nutrients



Effects of **parasympathetic stimulation**

- **Gastrointestinal system**: increases motility and secretory activity.
- **Glands**: increases secretory activity (but remember sweat glands are under sympathetic control).
- **Heart**: decrease rate of contraction (bradycardia).
- **Pupil**: control pupil diameter by papillary light reflex (myosis) (regulates the amount of light falling on retina).
- Accommodation of the **lens** for near vision. → by changing convexity of the lens by contraction of some smooth muscles there.
- Voiding the **urinary bladder** (micturation).

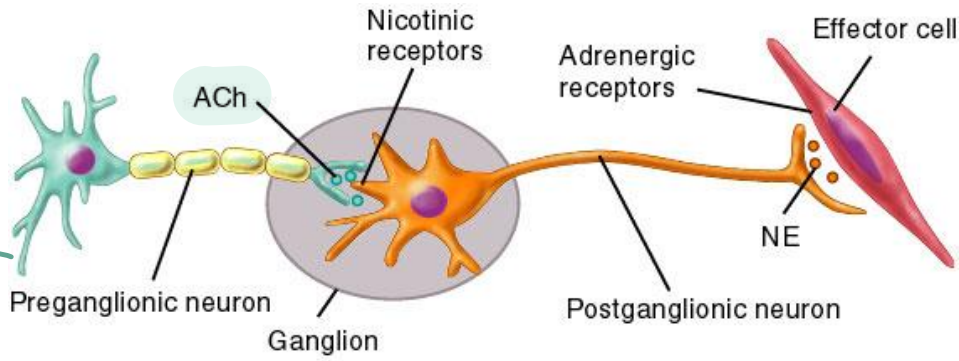
MOLECULAR BASIS OF PHYSIOLOGICAL ACTIONS OF THE ANS

- neurotransmitters
- types of receptors



Fig. 17.06

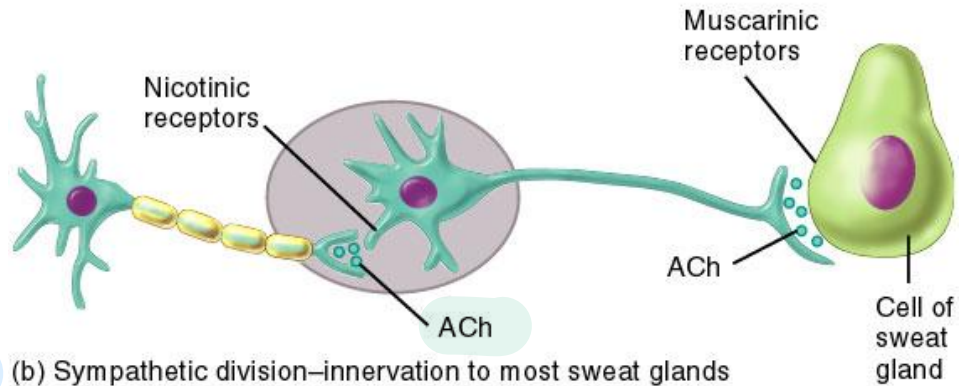
• the Preganglionic fibers are releasing ACh as neurotransmitters in both divisions (sympathetic + Para.)



(a) Sympathetic division—innervation to most effector tissues

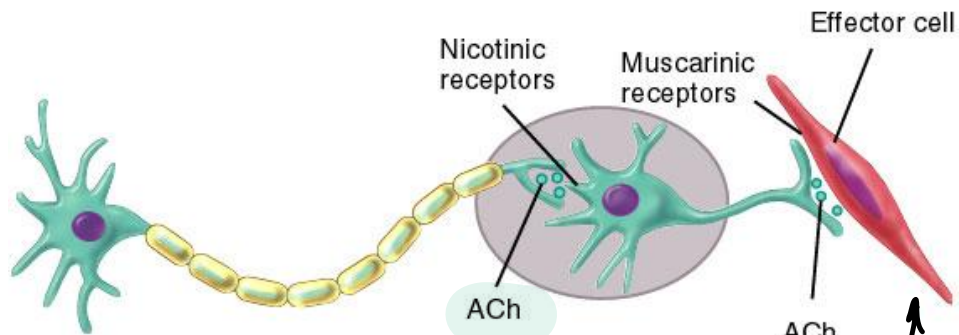
1

• Postganglionic
→ Parasympathetic Acetylcholine
→ Sympathetic norepinephrine
↓
exception



(b) Sympathetic division—innervation to most sweat glands

2



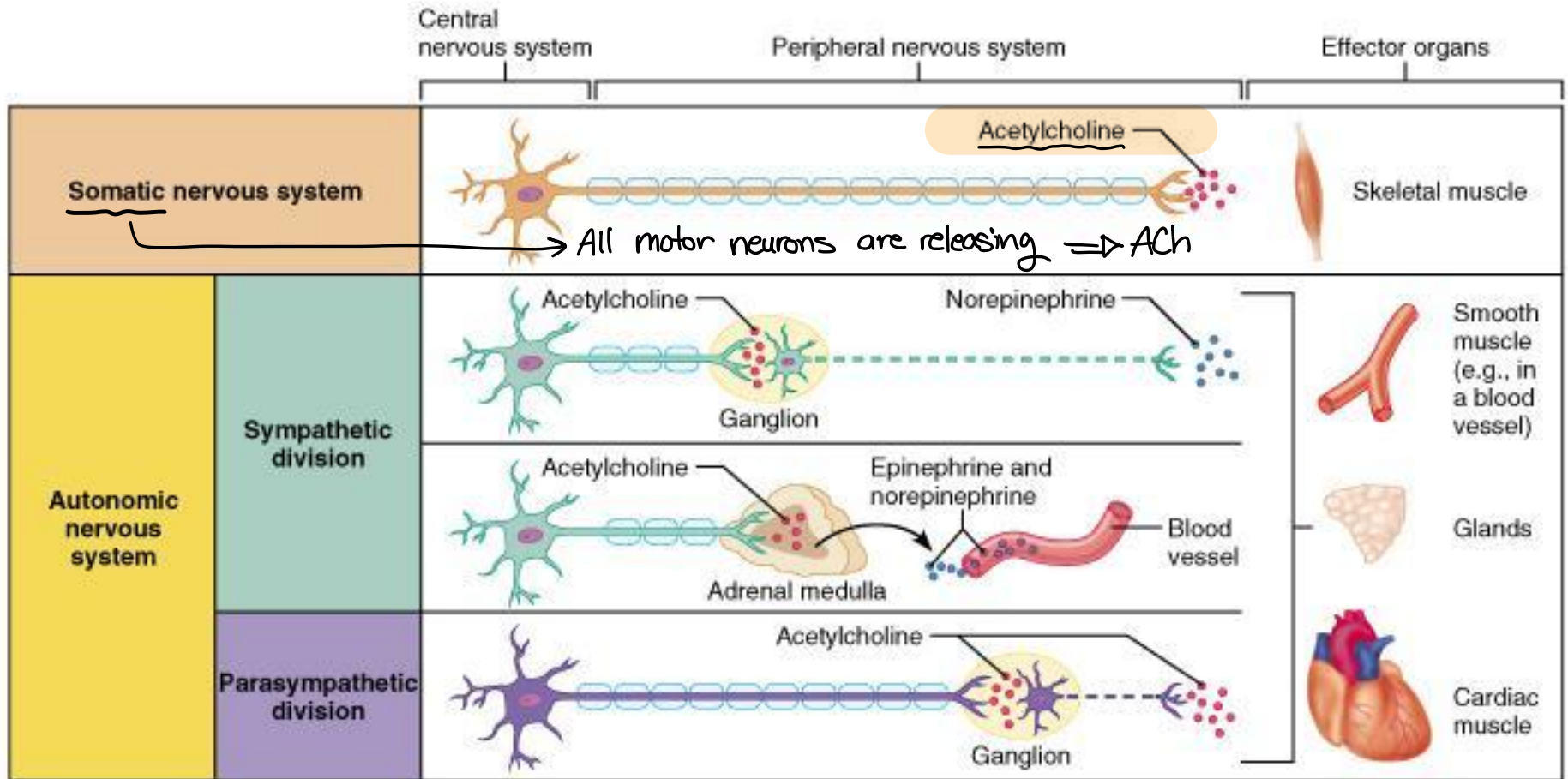
(c) Parasympathetic division

3

↳ effector: Sweat glands / pilomotor muscles = Acetylcholine



difference Between Somatic + Autonomic



Key:

- = Preganglionic axons (sympathetic)
- - - = Postganglionic axons (sympathetic)
- ⊖ = Myelination
- = Preganglionic axons (parasympathetic)
- - - = Postganglionic axons (parasympathetic)

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– Neurotransmitters

- At ganglion: preganglionic neurons of both sympathetic and parasympathetic release **acetylcholine (ACh)**.
- Effector organs:
 - parasympathetic fibers release **acetylcholine**
 - Sympathetic: **norepinephrine**.
- An exception for sympathetic nerves to sweat glands, which release **acetylcholine (ACh)**.

Don't forget !!
- neurons synapsing with endocrine cells of glands → Are 1st neurons of sympathetic division ⇒ releases ACh



Receptors and Signal transduction mechanisms

- At ganglia: sympathetic and parasympathetic have nicotinic receptors at the post synaptic membrane
↳ nicotine can stimulate them
↳ Muscarine can activate them.
- on effector cells: Muscarinic receptors.

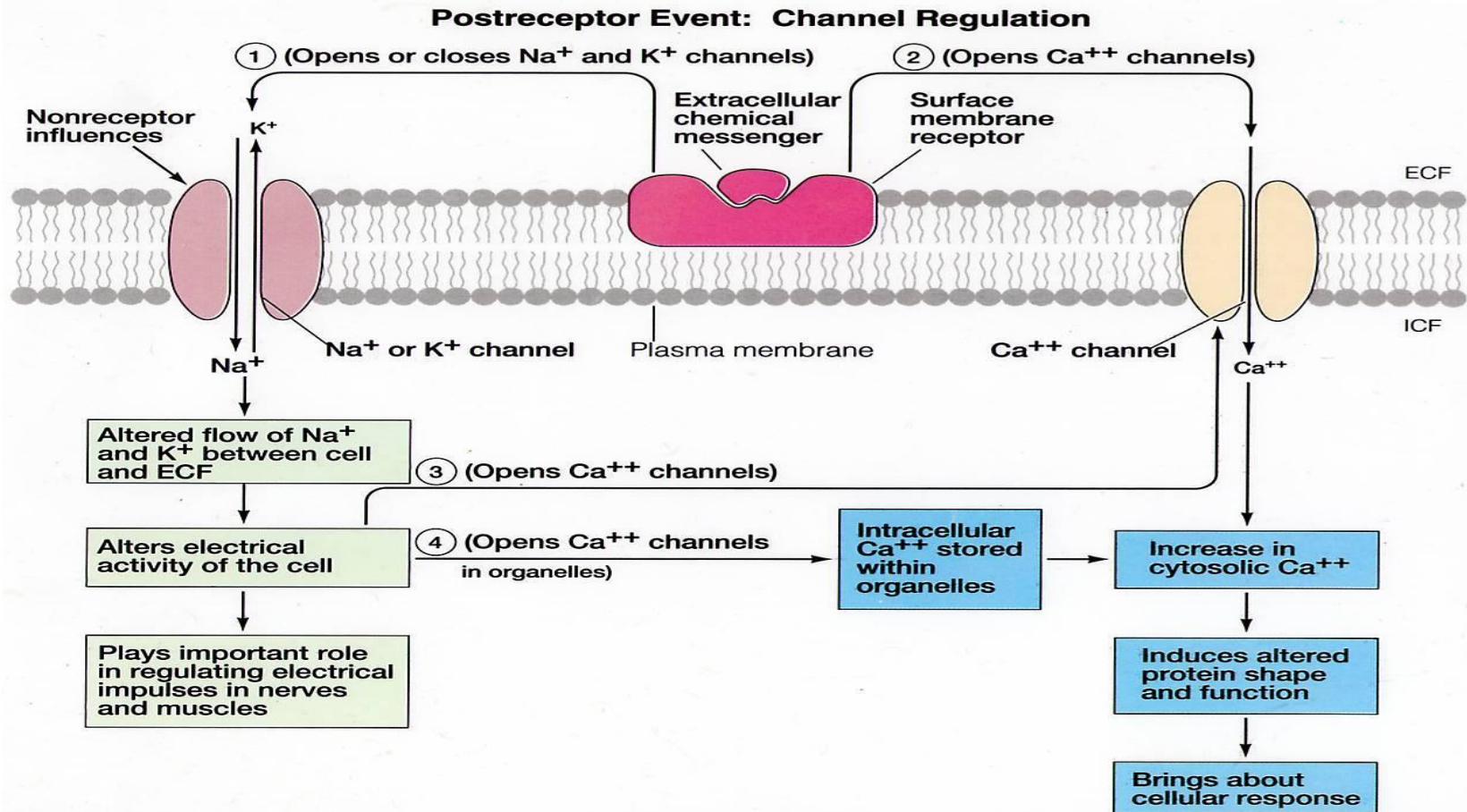
* ACh is also released by terminals of Motor Neurons of the Somatic System → at the level of skeletal muscles → type of receptors of ACh are called nicotinic receptors

* At which plant we can find Muscarine ?

Some toxic Mushroom is rich with Muscarine → so if somebody has ingested that mushroom they will develop or activate all muscarinic receptors in the body.



Signal Transduction Mechanism for Nicotinic receptors: Activation of Na⁺ Channels



- At the level of ganglia \rightarrow nicotinic receptors are linked to chemical gated Na^+ channels

\rightarrow Signal Transduction Mechanism :

- once we have activated the receptors
- Activation of chemical gated Na^+ channels
- Depolarizing membrane
- AP at the 2nd neuron.

Receptors and Signal transduction mechanisms

Muscarinic Receptors (M1-M5)

Inhibitory:

Conductive tissue of heart

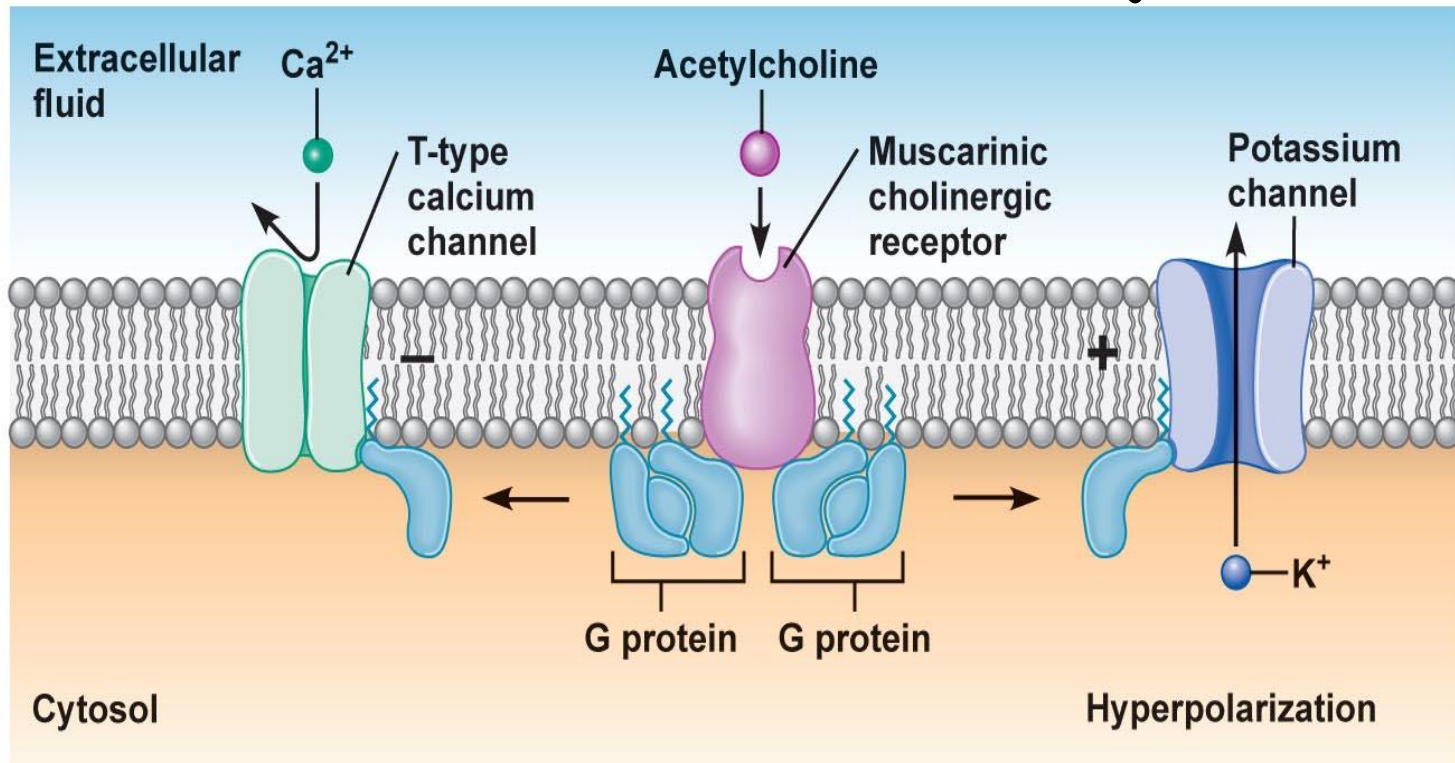


M2 in the heart: G protein \rightarrow K⁺ channel \rightarrow slow
the rate of depolarization. <sub>↓
reduce</sub>



inactivates
↓
Gi → adenylate Cyclase → reduces

cAMP → increasing Activity of K^+ channels
+ Decreasing // of T-type of Ca^{2+}
and Na^+ Channels.



(b) Parasympathetic



Receptors and Signal transduction mechanisms

Muscarinic Receptors (M1-M5)

Excitatory Receptors: (M1, M3, M5)

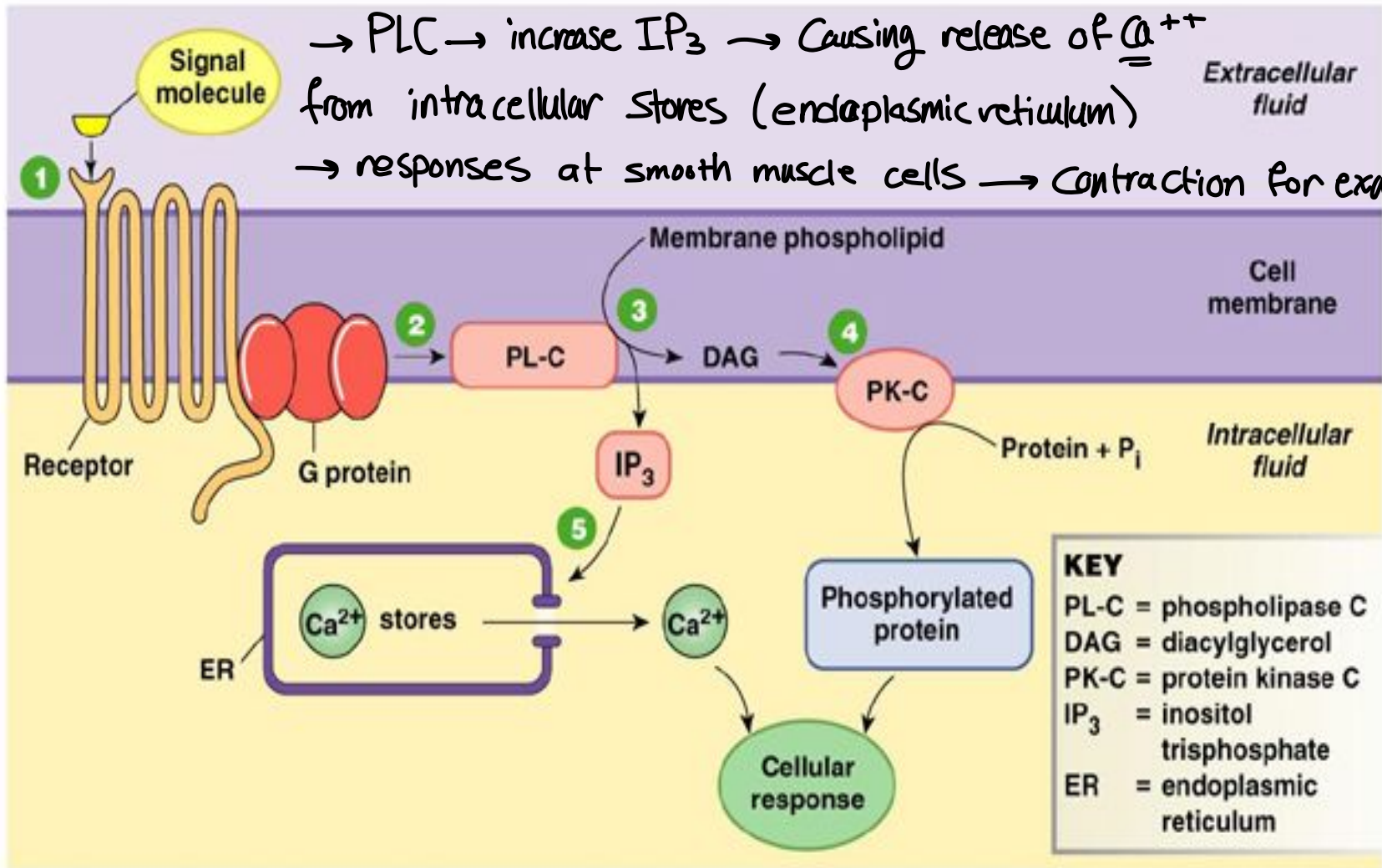
Found on **smooth muscle and glands** are coupled

Gq protein \rightarrow phospholipase C $\xrightarrow{\text{GIT}}$ $\xrightarrow{\text{Activate.}}$

This enzyme increases production of inositol-1,4,5-trisphosphate (IP3)



→ PLC → increase IP_3 → Ca²⁺ from intracellular stores (endoplasmic reticulum)
 → responses at smooth muscle cells → contraction for example.



- 1 Signal molecule activates receptor and associated G protein.
- 2 G protein activates phospholipase C (PL-C), an amplifier enzyme.
- 3 PL-C converts membrane phospholipids into diacylglycerol (DAG), which remains in the membrane, and IP₃, which diffuses into the cytoplasm.
- 4 DAG activates protein kinase C (PK-C), which phosphorylates proteins.
- 5 IP₃ causes release of Ca²⁺ from organelles, creating a Ca²⁺ signal.



Activation of Muscarinic Receptors

During Muscarine

intoxication → Patients

who ingested toxic mushroom develop:

- Stimulation of secretory activity: salivation, tearing, sweating, nasal and bronchial secretion.
- Increase gastrointestinal tract motility → vomiting and diarrhea.
- Contraction of urinary bladder → urination.
- Slowing of the heart → Bradycardia.



to reverse effects of Muscarinic intoxication → Drug ↙

Blocking of Muscarinic Receptors by ATROPIN

Signs related to giving high dose of Atropin:-

- Inhibition of glandular secretions → dry mouth, dry eyes, and dry nasal passages.
- Tachycardia. (increase heart rate).
- Loss of pupillary light reflex → Mydriasis
- Loss of ability to focus the lens for near vision.

→ you are following the heart rate of patient + pupillary light reflex
once you have any increase in the heart rate → you can stop giving Atropin



Receptors and Signal transduction mechanisms

Adrenergic receptors:

These receptors respond to **catecholamines:** (epinephrine (EP) and norepinephrine (NE)).



Receptors and Signal transduction mechanisms

• Adrenergic Receptors

Alpha receptors:

- **The alpha 1 (α_1)**: Excitatory: PLC \rightarrow IP3

\rightarrow Smooth muscle cells of vessels + Arterioles \rightarrow by activation \rightarrow Vasoconstriction

- **Alpha2** receptors: Nerve Adrenergic terminals \rightarrow reduce NE release

Alpha 2 Heteroreceptors: Nonadrenergic -

Gi \rightarrow Adenylate cyclase \rightarrow decrease cAMP



Receptors and Signal transduction mechanisms

Beta receptors:

- **Beta 1** (β_1) receptors: found on heart

↳ excitatory

- **Beta 2** (β_2) receptors: found on tracheal and bronchial smooth muscle, in the gastrointestinal tract, and on smooth muscles of blood vessels supplying skeletal muscles

↳ Inhibitory

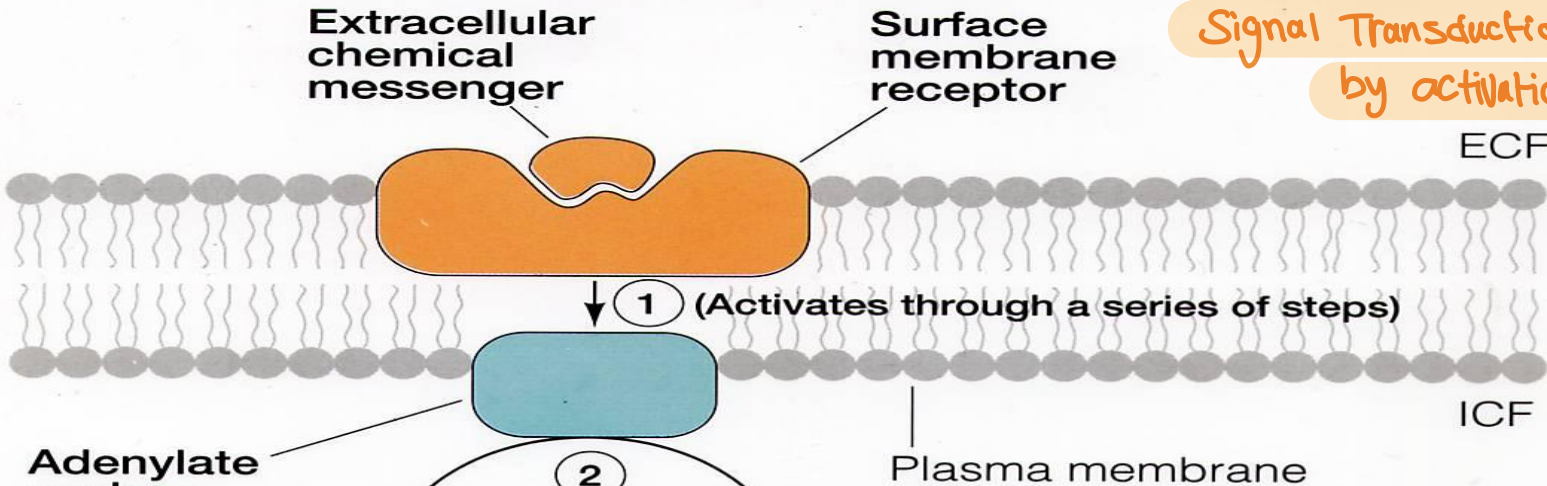
Gs → Adenylate cyclase → increase cAMP

↳ Both B_1/B_2 work by activation of AC → Heart → increases heart rate
↳ Smooth Muscles → reduce relaxation of smooth muscles.

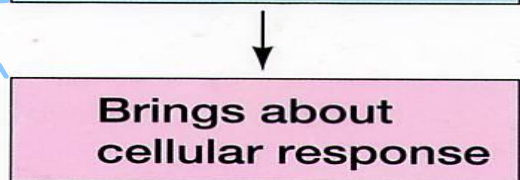
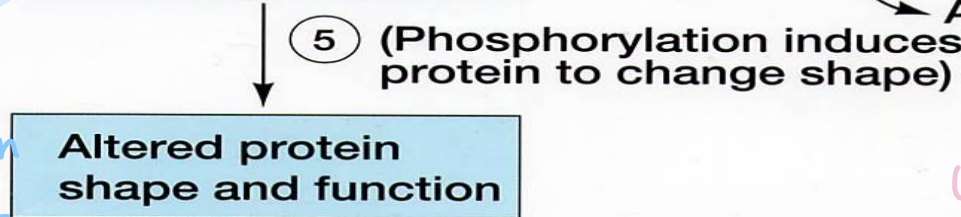
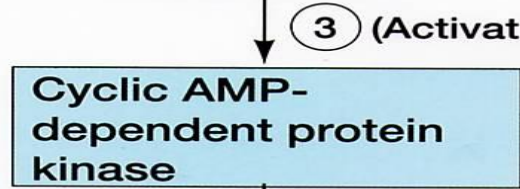


Postreceptor Event: Cyclic AMP Second Messenger System

Signal Transduction Mechanism
by activation of Beta
receptors.



increase in c-AMP
may cause
excitation
inhibition




↑ c-AMP

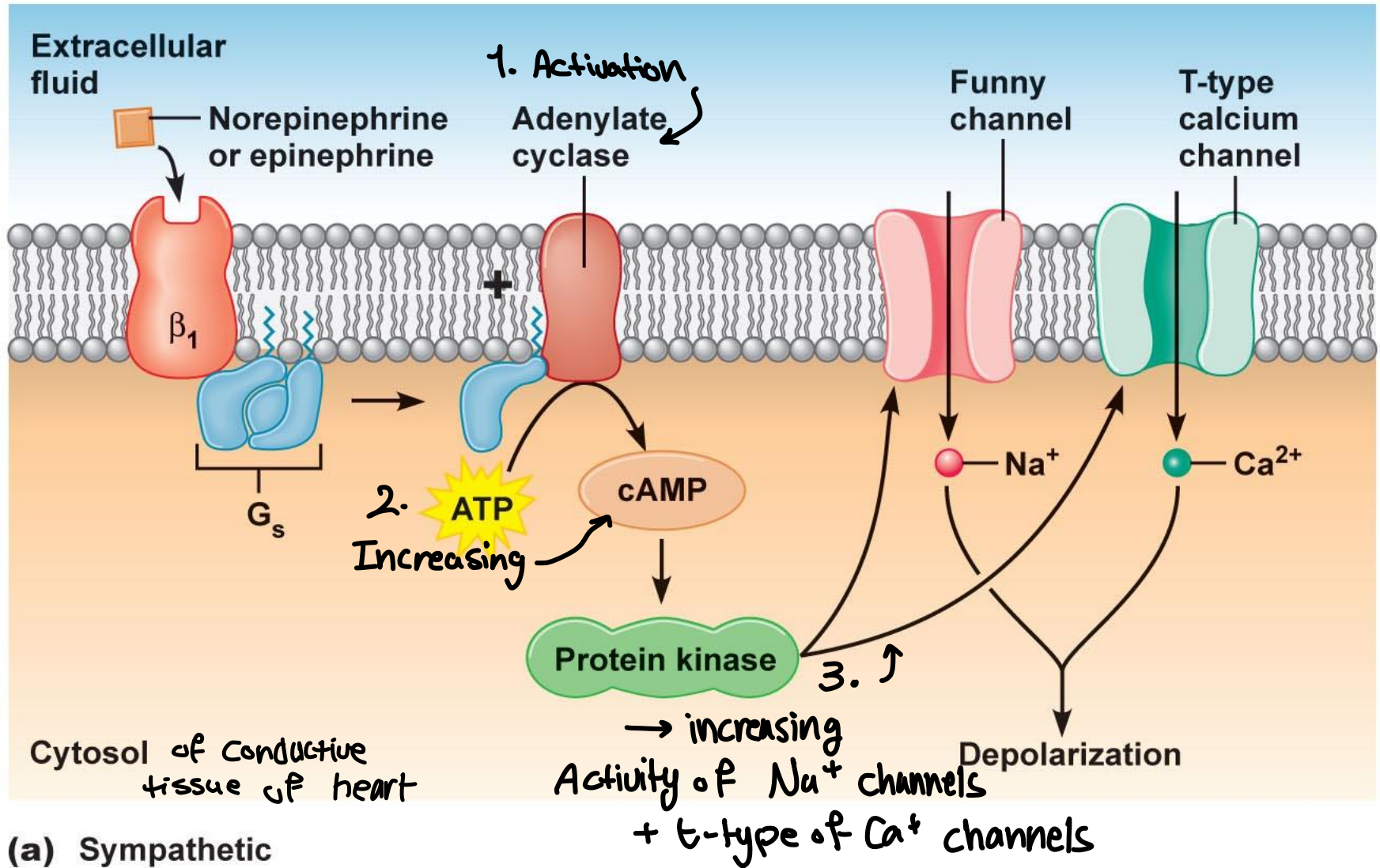
Heart
↓
increases Heart Rate

Smooth Muscles
↓
Reducing Contraction activity
+
Inhibition of SM cells

depending on type of cells where we have Beta receptors.



What happens at the conductive tissue of the Heart



(a) Sympathetic

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→ Decrease in K⁺ channels



Summary

Muscarinic receptors: Activated by Ach.

- Inhibitory: **M2, M4**
- Excitatory: M1, M3, M5

Adrenergic receptors: activated by Epinephrine and Norepinephrine.

- Alpha (α) Receptors: α_1 , **α_2 , α_2 hetero-**
- Beta (β) Receptors: β_1 , **β_2** , ... β_3

- **Beta 2 (β_2)** receptors: found on tracheal



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GOOD LUCK

