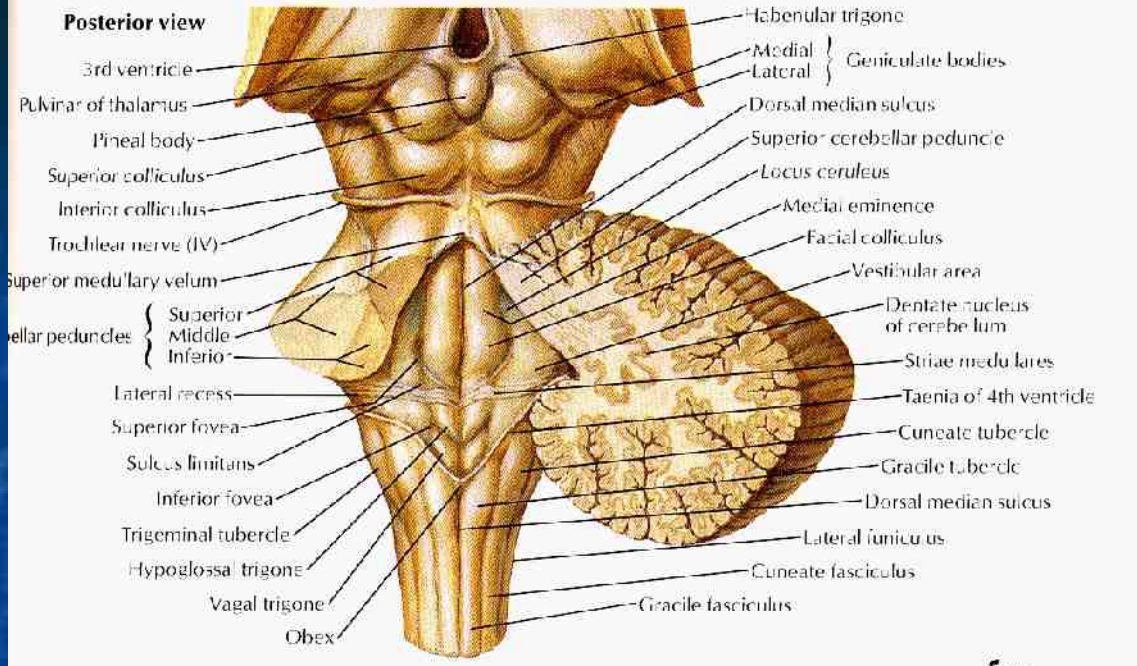
THE CEREBELLUM

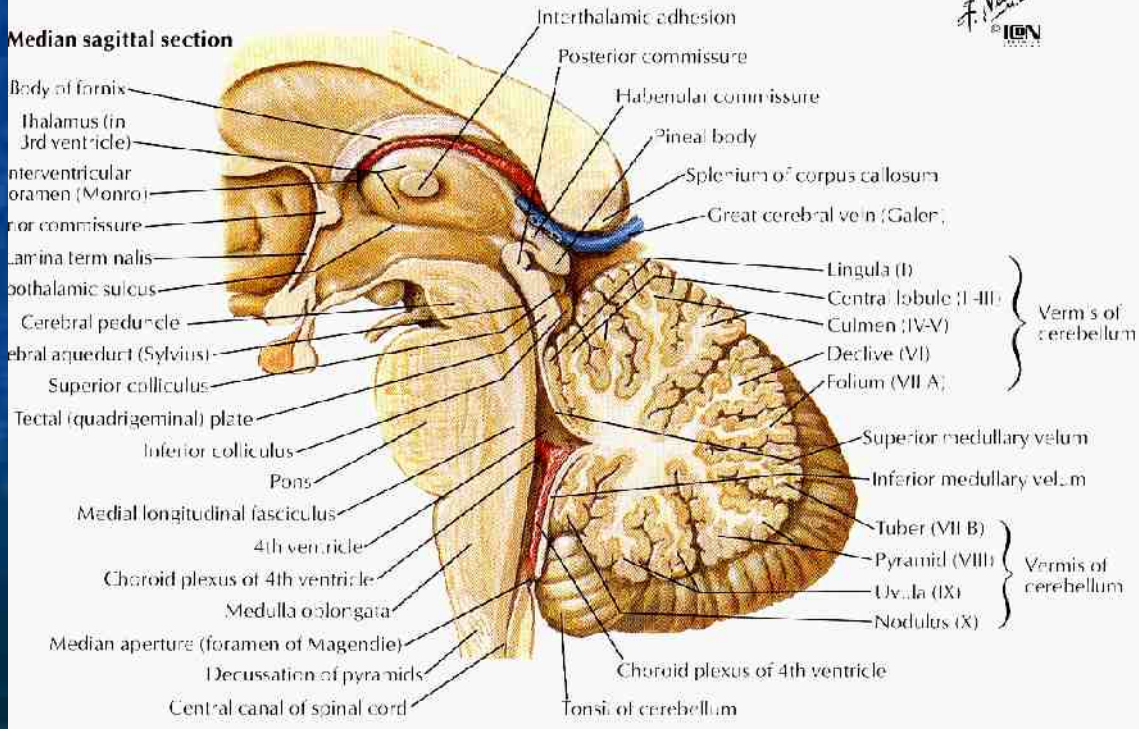
Institute of Anatomy, 2nd and 1st Medical Faculty
R. Druga

MOZEČEK

Anatomický ústav 2.a 1. LF
R. Druga

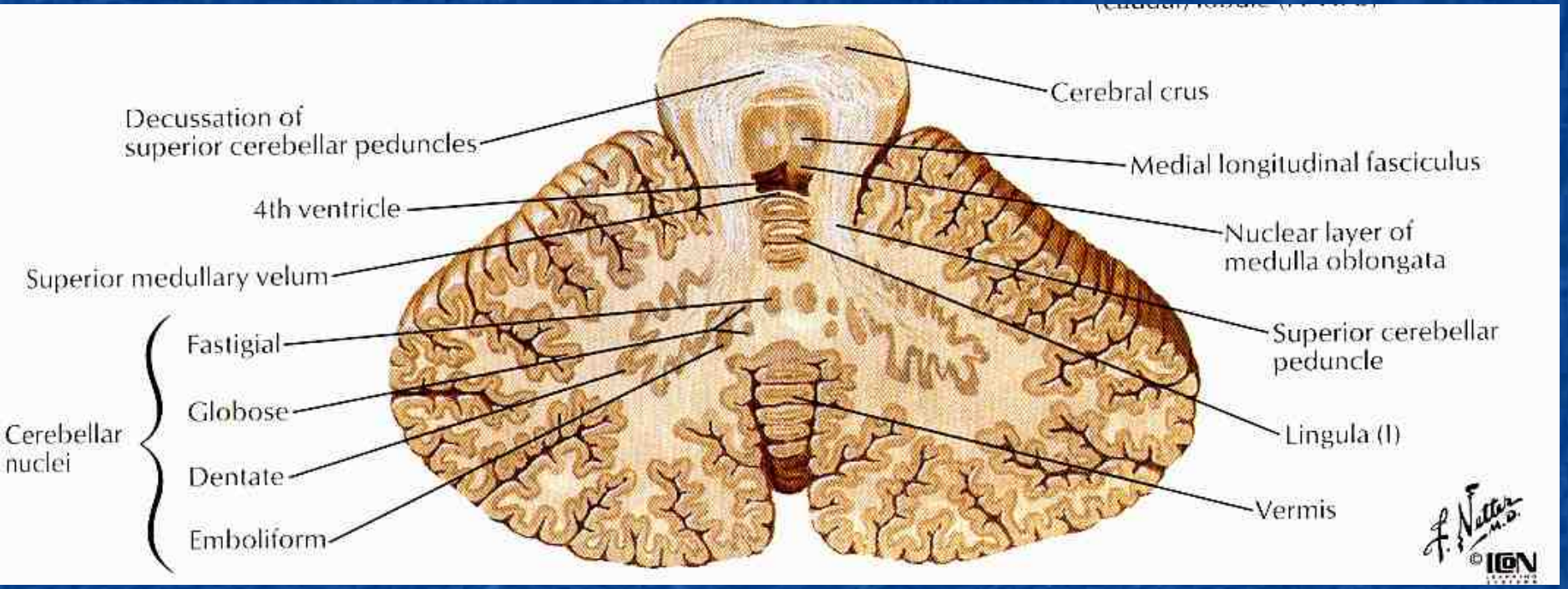
„Little brain „





Cerebellar gyri = folia

Cortex, subcortical white matter, nuclei, Kůra, bílá hmota, jádra

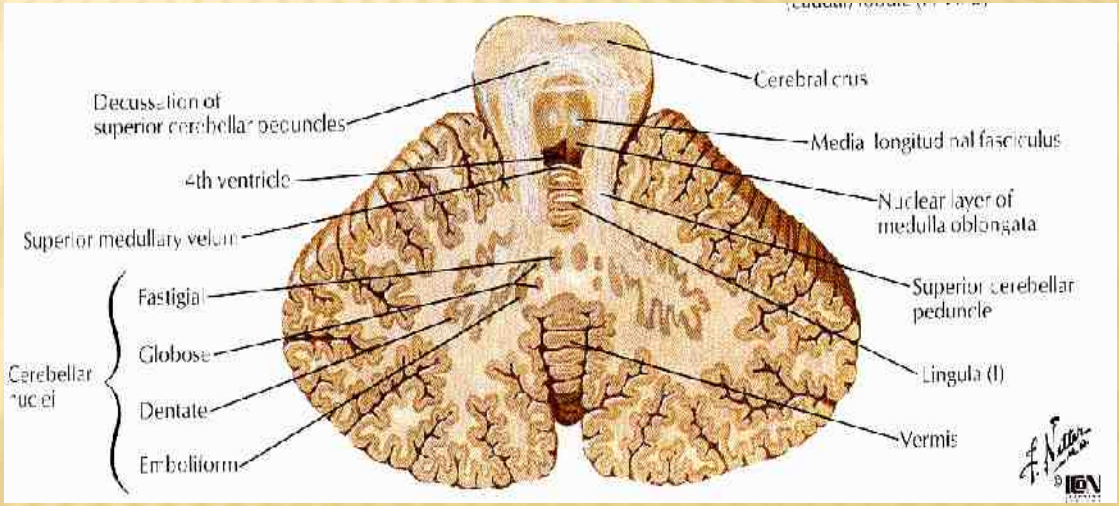


Excessive folding of the cerebellar surface (cortex)

Mimořádně bohatá gyrifikace kůry

THE CEREBELLUM – RELATIONS AND STRUCTURE

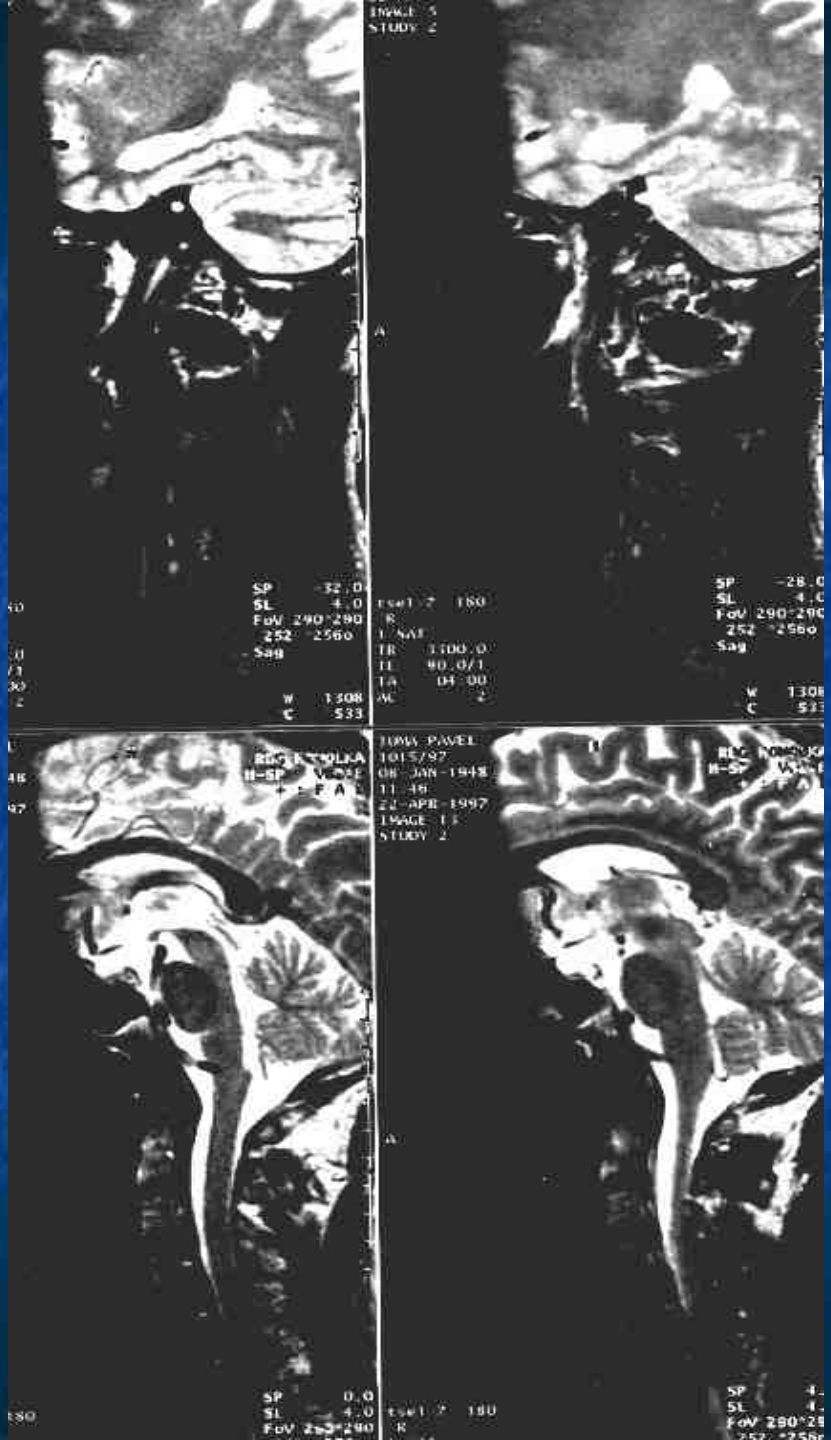
- Little brain
- Located in the posterior crerebral fossa
- Connected with the brain stem- by peduncles (inferior, middle, superior)
- Is covered by the cerebellar cortex (3 layers). Cortex is extensively folded (folia-oriented mediolaterally)
- White matter fibers going to and coming from the cortex
- In the white matter are the cerebellar nuclei



MR examination

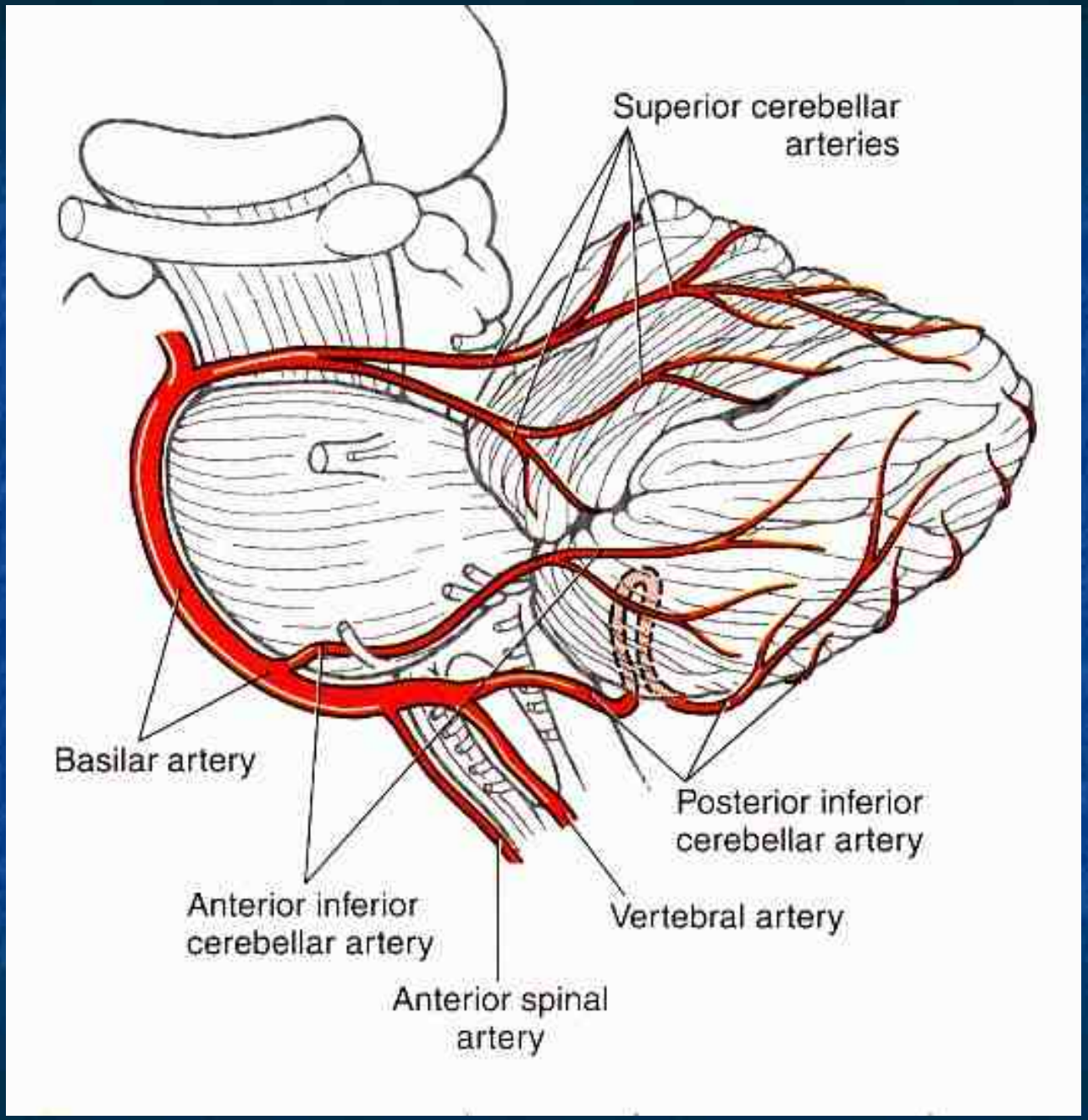
Fossa cranii posterior

Posterior cranial fossa



Arterial supply

SCA



AICA

PICA

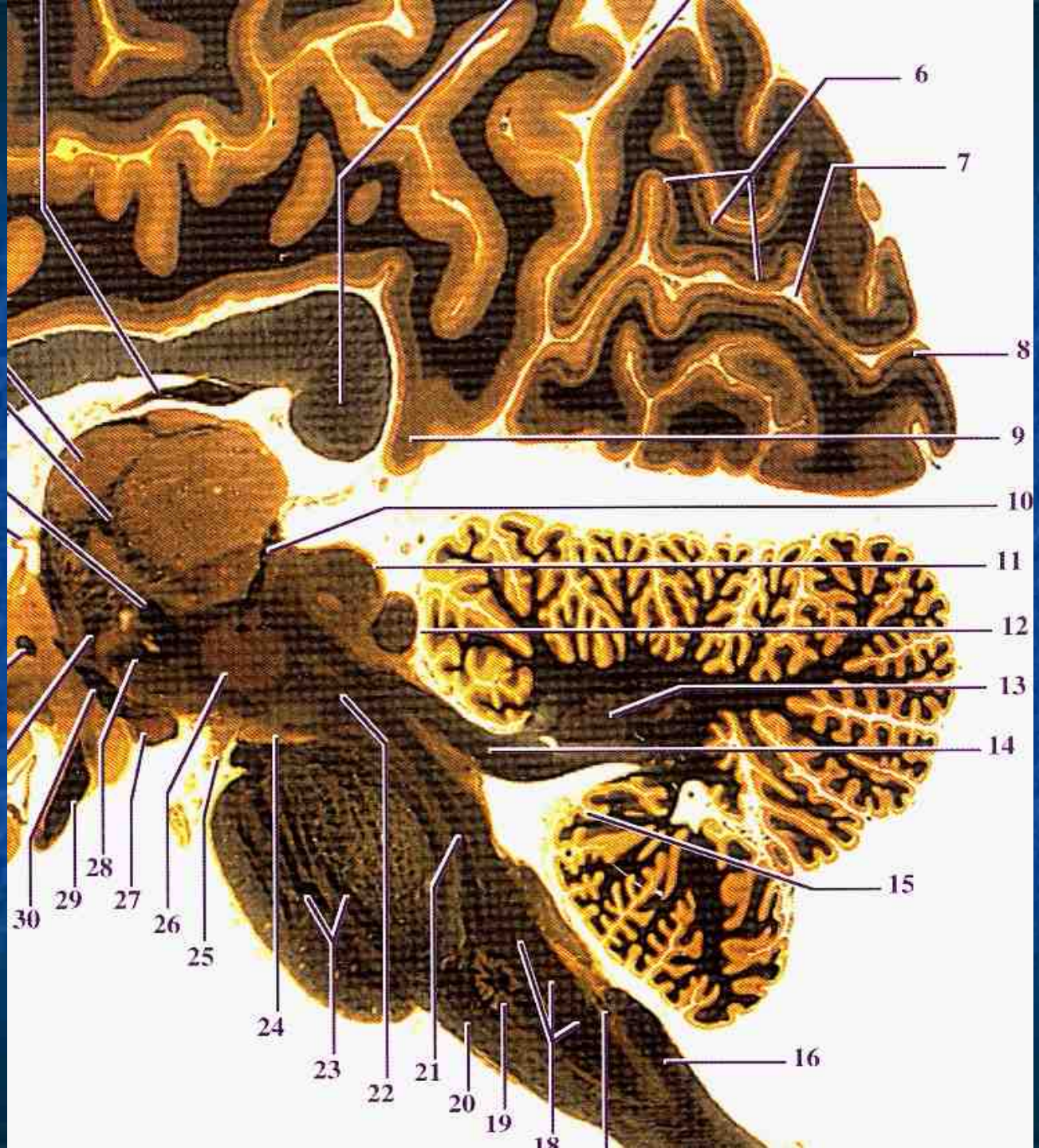
Weigert Staining

Myelinizovaná vlákna

White matter has a treelike appearance =

arbor vitae

(„tree of life „)

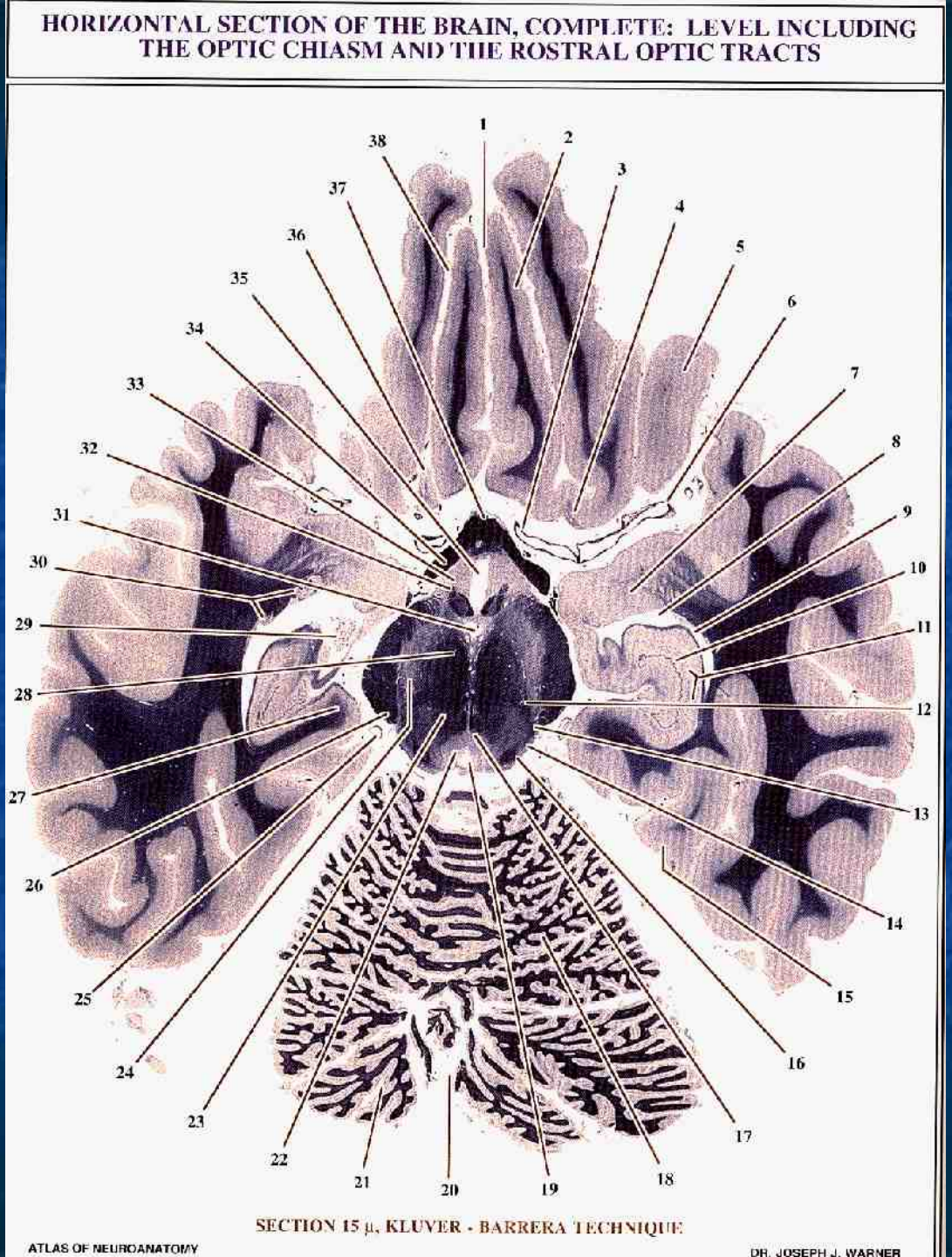


Klüver - Barrera

staining

Folia are not interupted

Folia jsou nepřerušena



Cerebellar hemispheres

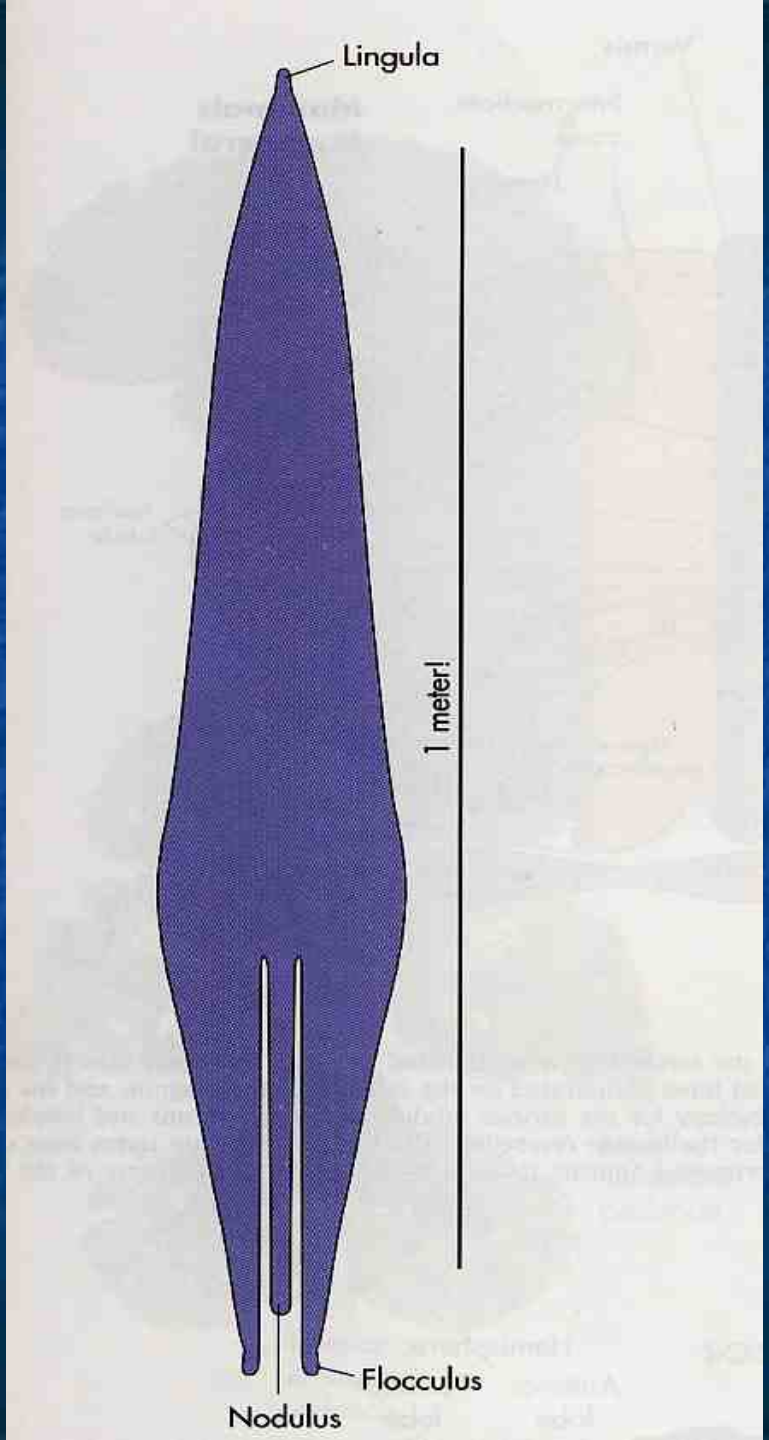
and vermis

Unfolded surface of the cerebellum

Rozvinutá kůra do plochy (dvourozměrná rekonstrukce)

Breadth = 10 – 15 cm

Šířka = 10 – 15 cm
Délka = 1 mm



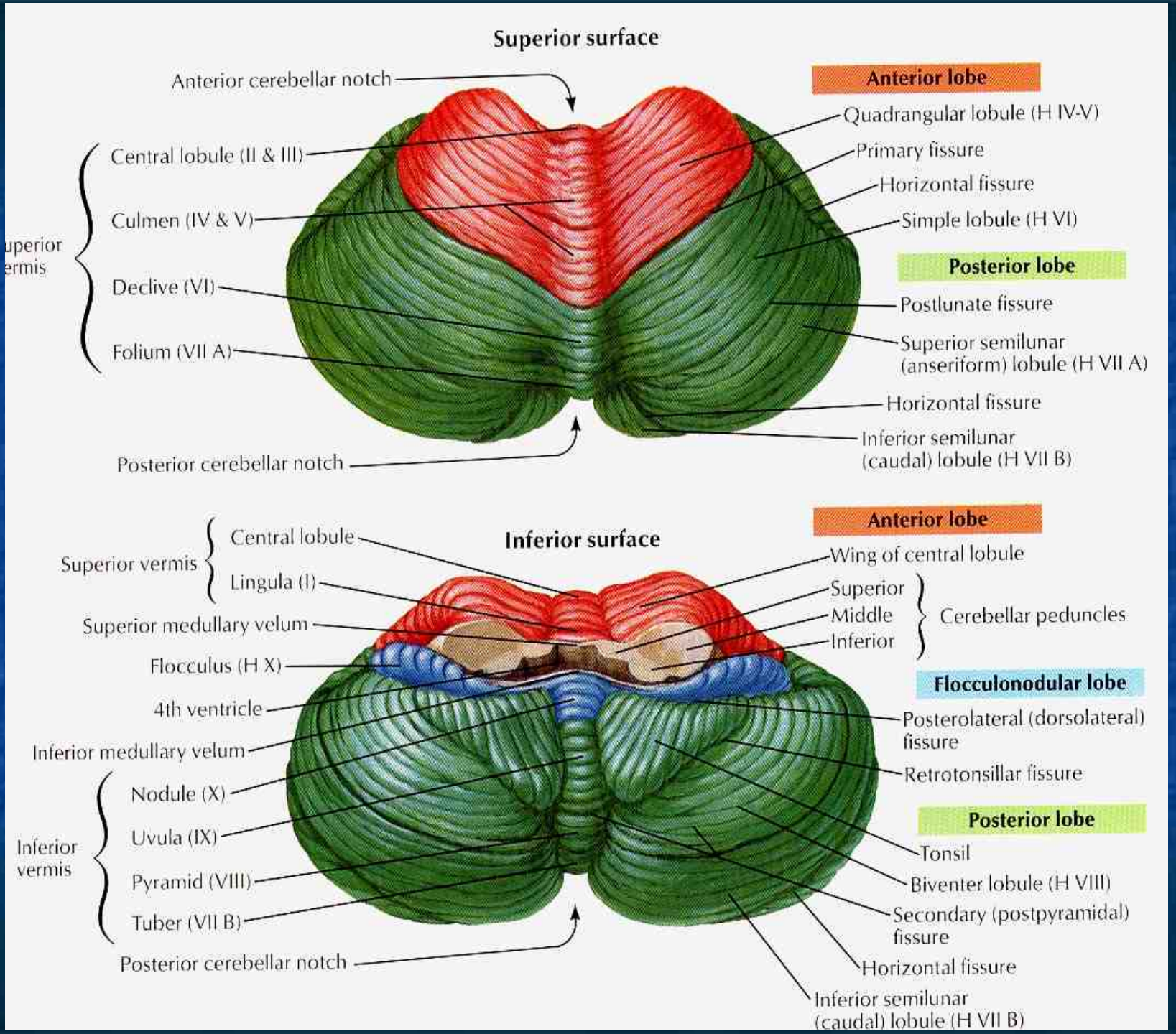
THREE CEREBELLAR LOBES

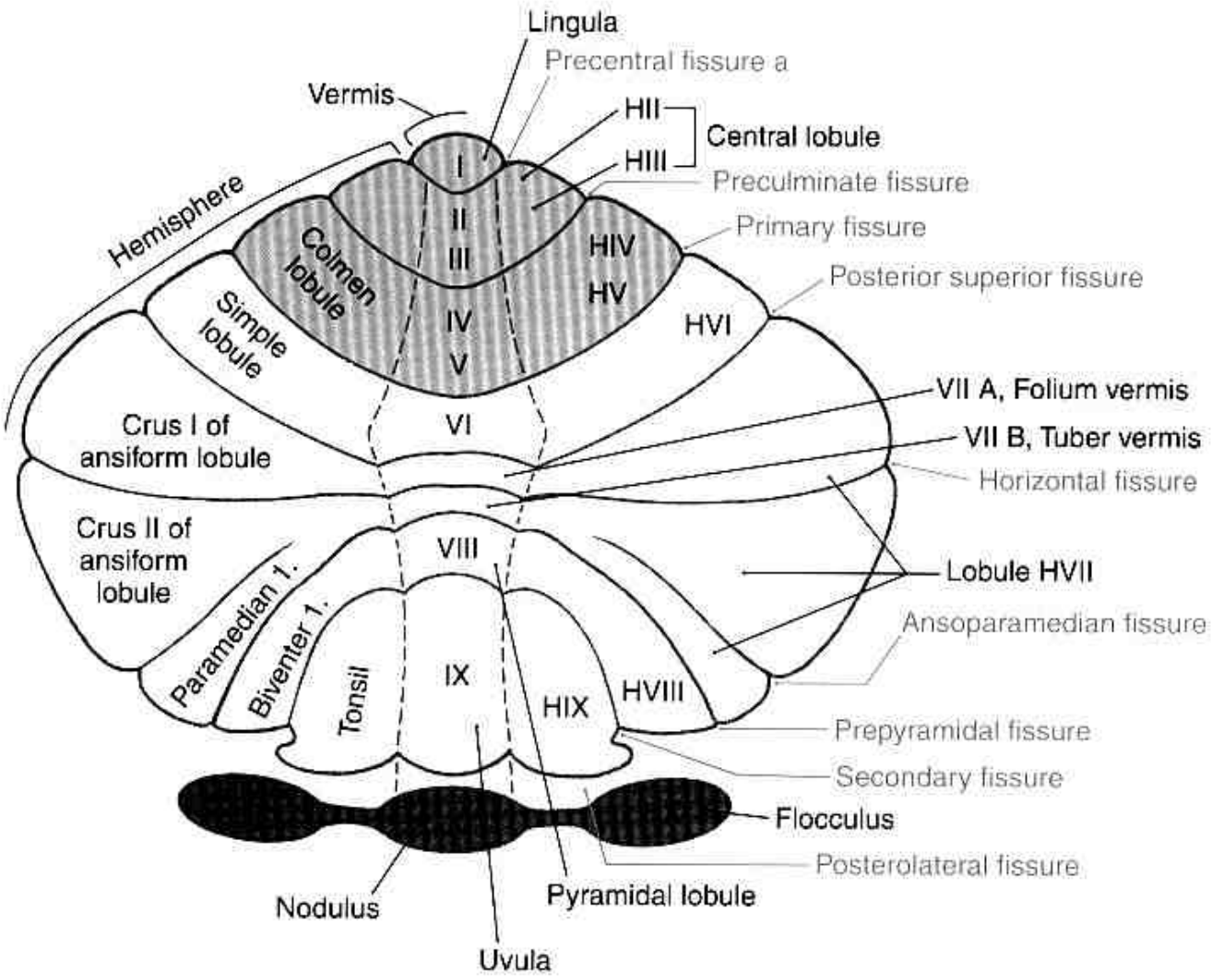
- Anterior lobe (I – V lobules)
- Posterior lobe (VI – IX lobules)
- Flocculo-nodular lobe (lobule X)

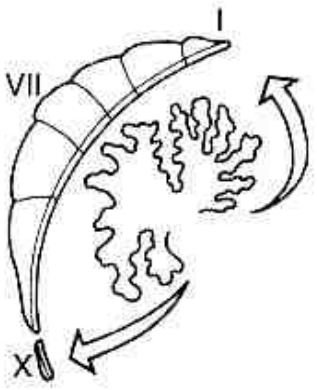
Deep

Fissures

Fissury







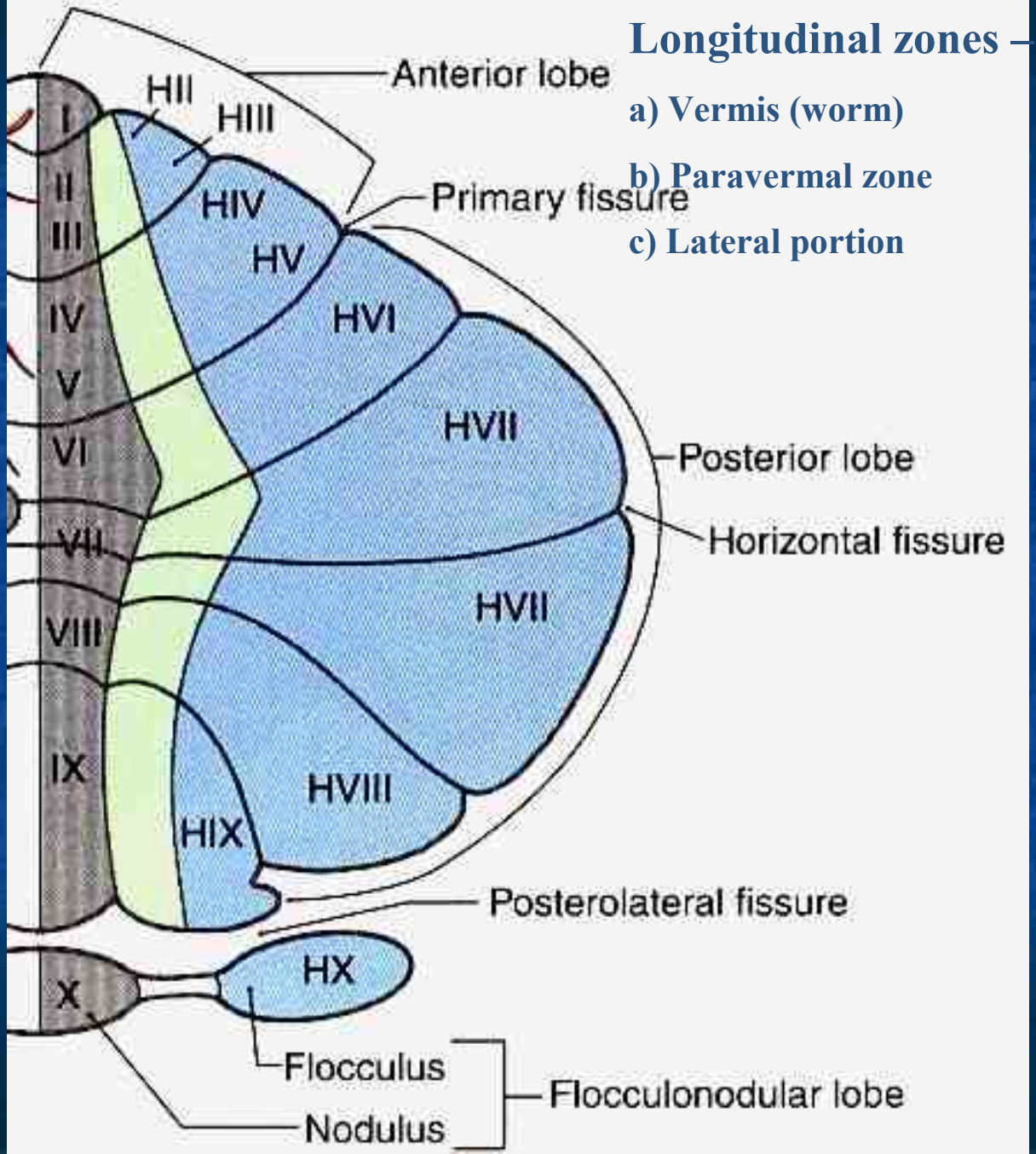
VII B – boundary between superior and inferior surface

Anterior lobe Posterior lobe Flocculonodular lobe

ure 27–2. Unfolded view (see *upper right*) of the cerebellar cortex showing lobes, lobules (by name and number), and main ures (printed in blue). The lobules of the hemisphere are designated by the prefix "H," to show which lobule of the isphere (H) is continuous with its corresponding (designated by the Roman numeral) vermal lobule. l., lobule.

Cerebellar lobes and Lobuli I – X

Lobi a Lobuli I - X

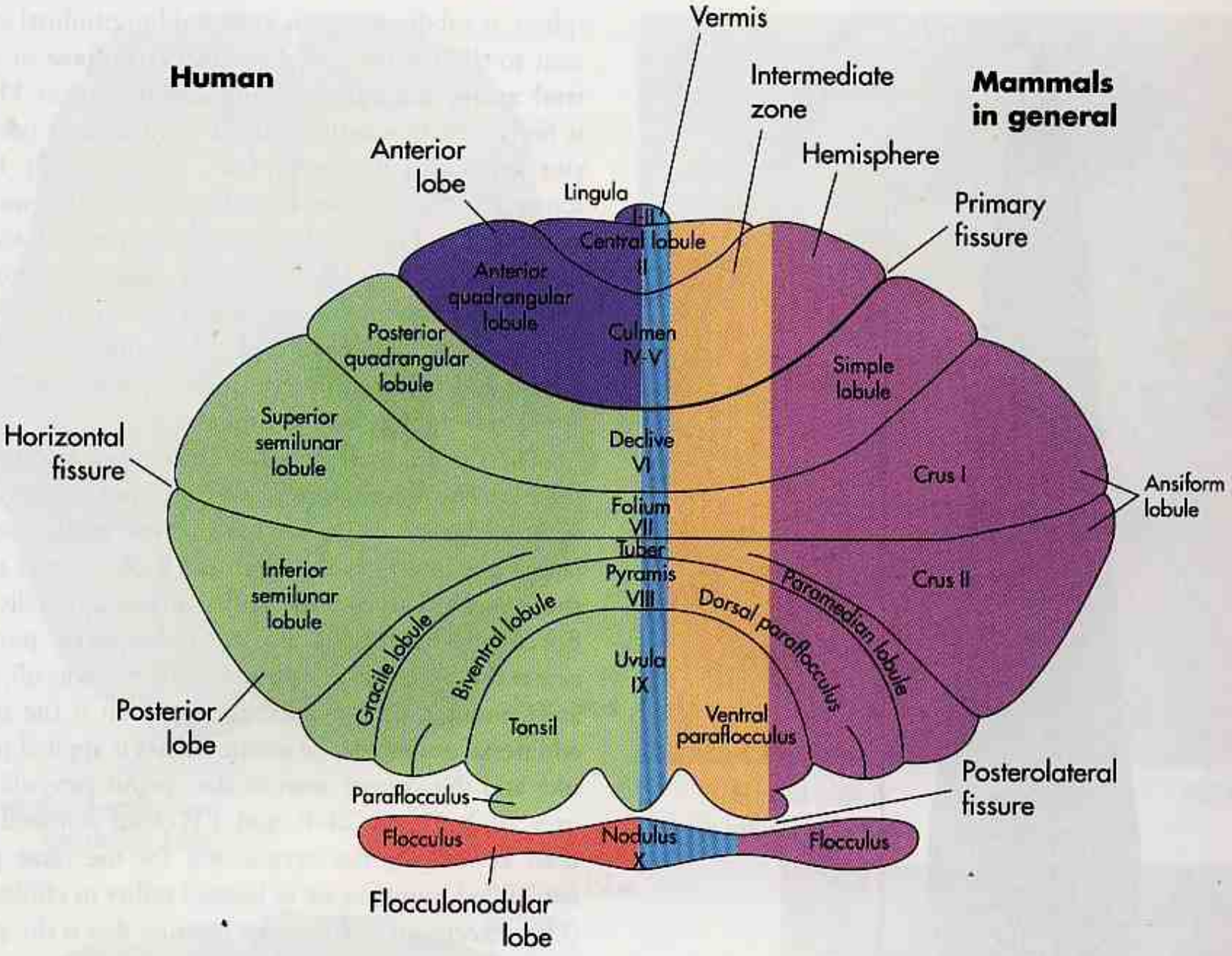


Longitudinal zones –

a) Vermis (worm)

b) Paravermal zone

c) Lateral portion


Human
Mammals in general
Vermis
Intermediate zone
Hemisphere
Anterior lobe
Lingula
I-II
Central lobule
II
Primary fissure
Anterior quadrangular lobule
Posterior quadrangular lobule
Culmen
IV-V
Simple lobule
Superior semilunar lobule
Horizontal fissure
Declive
VI
Crus I
Ansiform lobule
Folium
VII
Tuber
Inferior semilunar lobule
Pyramis
VIII
Crus II
Paramedian lobule
Dorsal paraflocculus
Gracile lobule
Biventral lobule
Uvula
IX
Posterior lobe
Tonsil
Ventral paraflocculus
Paraflocculus
Posterolateral fissure
Flocculus
Nodulus
X
Flocculus
Flocculonodular lobe

CEREBELLAR NUCLEI, neurons glutamatergic, excitatory, high spontaneous activity

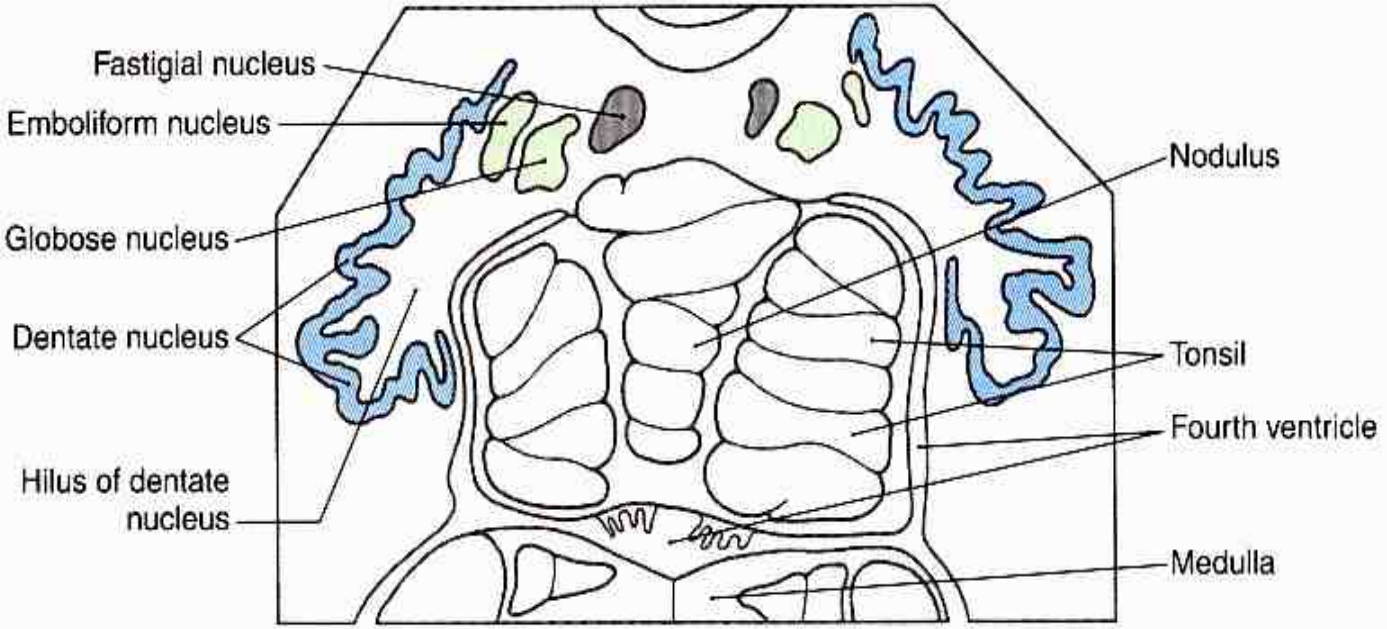


Figure 27–5. The cerebellar nuclei in cross section, drawn from a slide. The color coding of each nucleus corresponds to its appropriate zone in Figure 27–4.

Mozečková jádra, glutamátergní neurony, vysoká spontánní aktivita

STRUCTURE OF THE CEREBELLAR CORTEX - 3 LAYERS:

I. MOLECULAR LAYER - INHIBITORY INTERNEURONS

II. PURKYNĚ CELL LAYER – INHIBITORY PROJECTING NEURONS

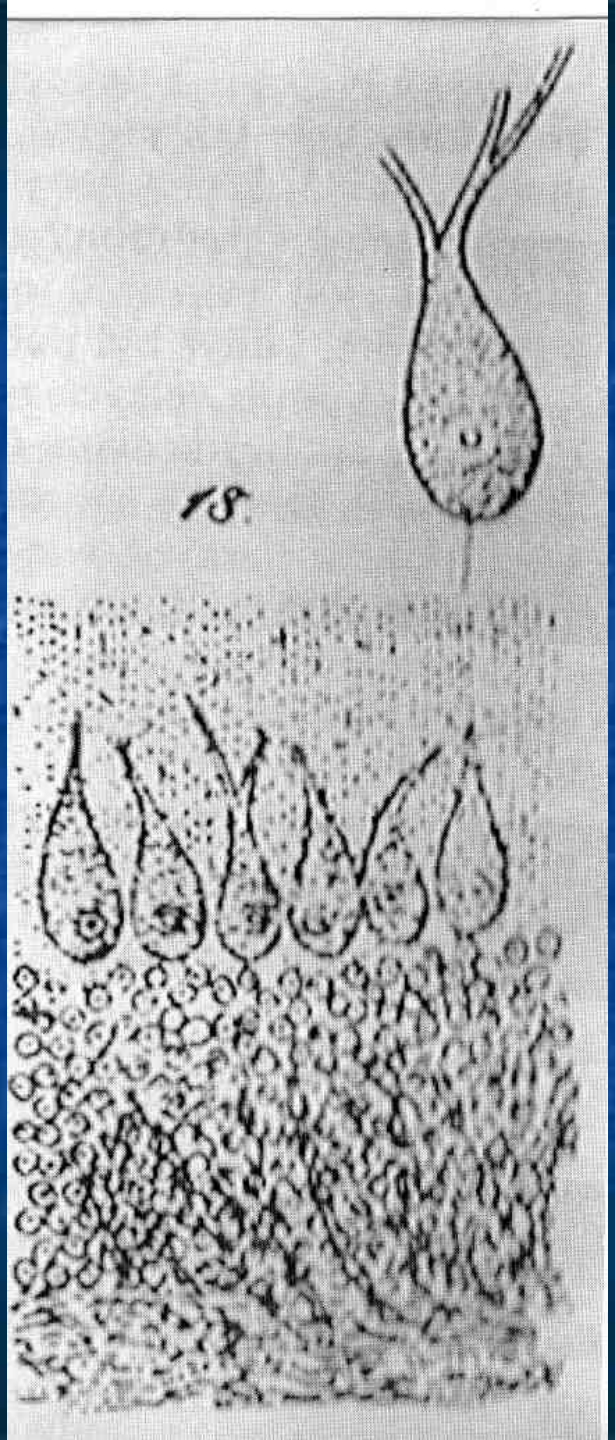
III. GRANULAR LAYER – PREVAIL EXCITATORY NEURONS

J.E. Purkyně 1837

Wroclaw, Poland

Breslau, Silesia

Purkyně ´s original lithography (engraving)



Purkyně observed for the first time large cells of the cerebellar cortex

R. y Cajal : Histologie du systeme nerveux, 1911.

Cellules de Purkinje. — Ces cellules, dont le nom rappelle celui du savant qui les a découvertes [1], sont volumineuses, ovoïdes, semi-lunaires ou mitrales et disposées en une rangée discontinue, juste aux confins de la couche plexiforme et de celle des grains. Leur diamètre, quelque peu variable suivant les mammifères, oscille chez l'homme entre 35 et 65 μ.

1. Purkinje, Bericht auf der Versammlung deutscher Naturforscher in Prag, 1837.

Santiago Ramon y Cajal 1911 (Cajal confirmed Purkyně´s findings)

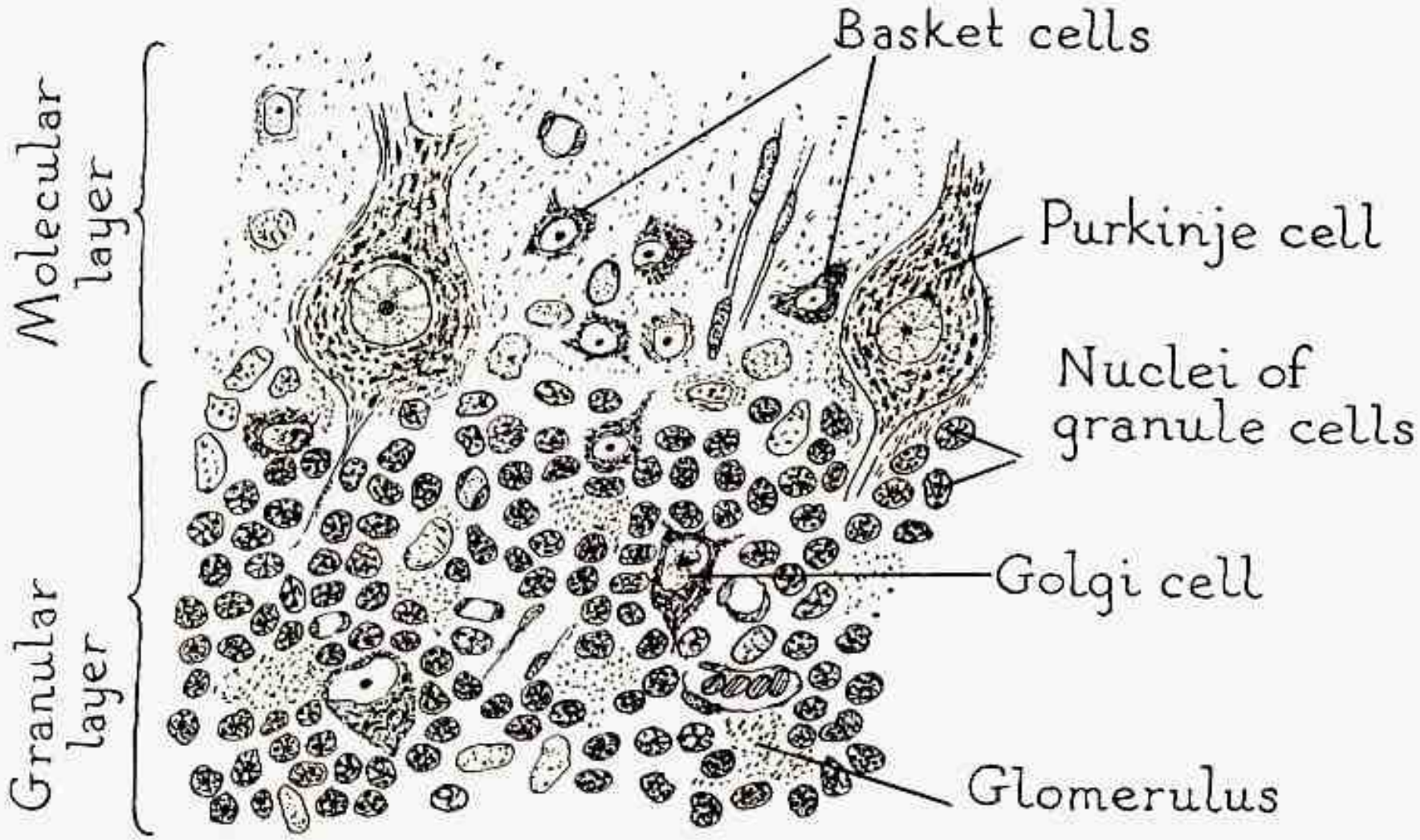


FIG. 246. Part of a section through human cerebellar cortex. Nissl stain. (After Cajal)

Basket cells, inhibitory interneurons, GABAergic

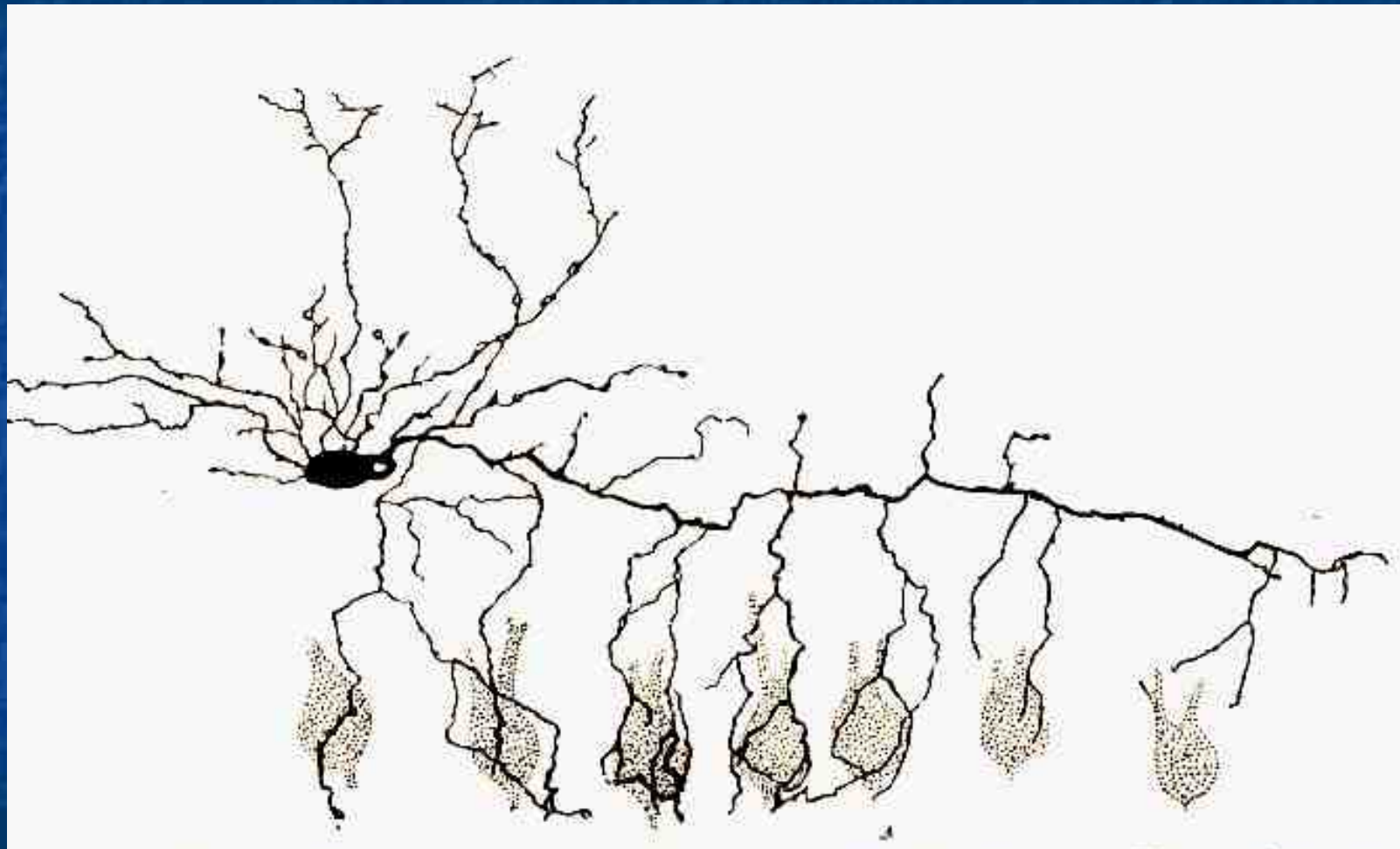


Abb. 265. Korbzelle mit ihrem Neuriten und dessen kollateralen Verzweigungen an den Purkinjeschen Zellen. Mensch. Golgi-Präparat. (Nach Jakob, Hdb. mikr. Anat. d. Menschen 4, I (1928))

Kompletní dendritická arborizace Purkyňovy b.

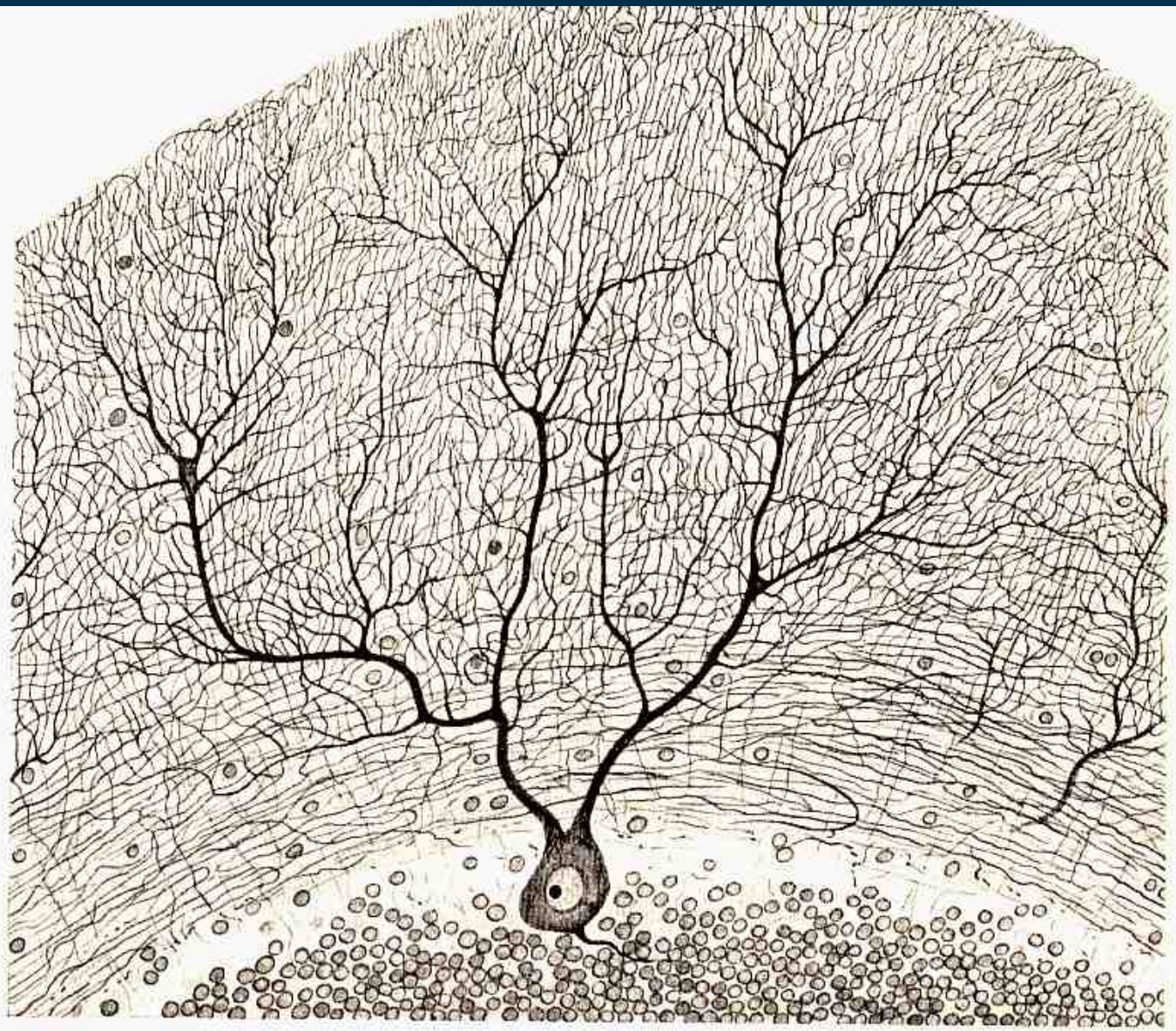
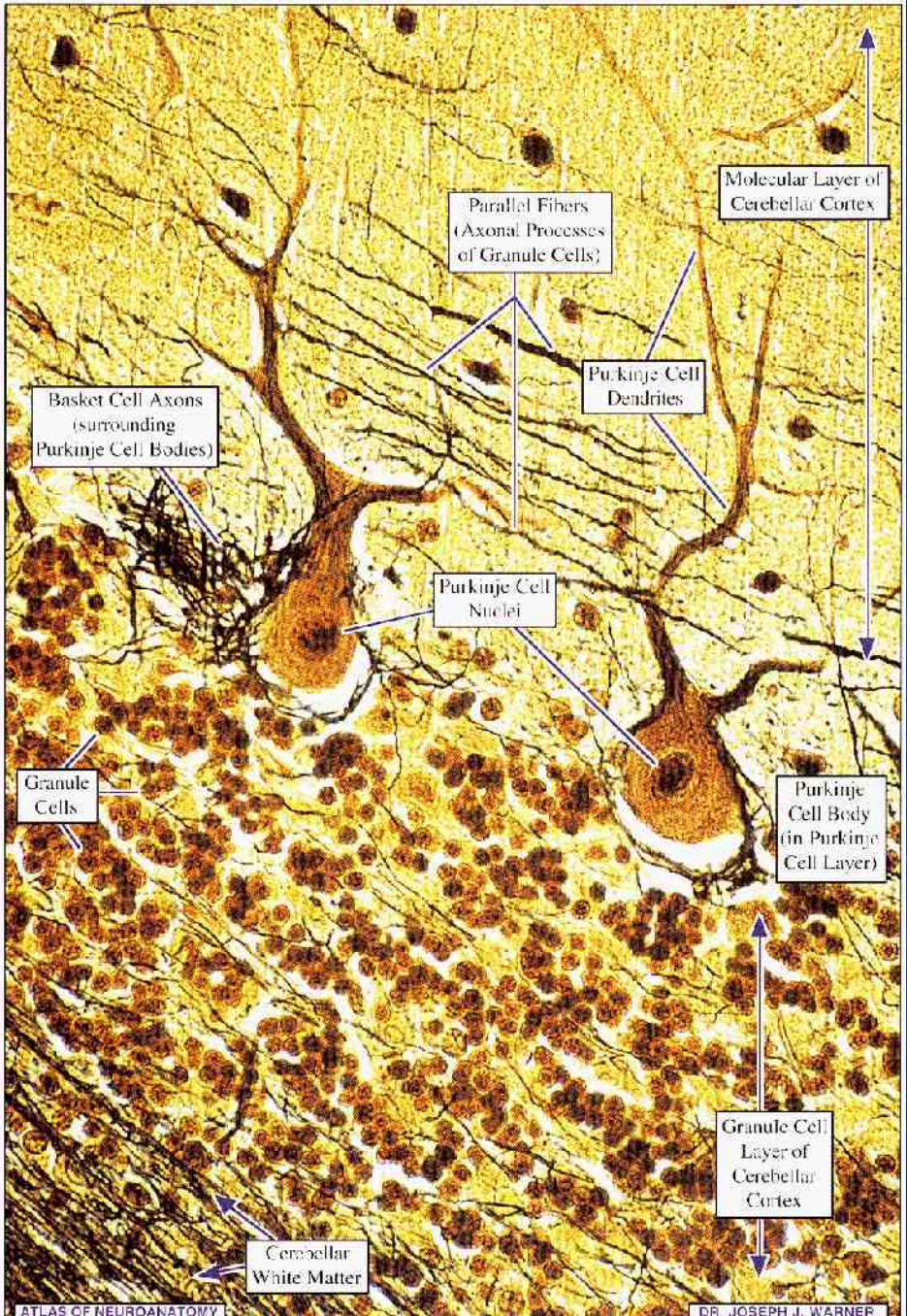


Abb. 261. Purkinjesche Zelle aus der menschlichen Kleinhirnrinde. Die Dendriten bilden untereinander ein zusammenhängendes Netzwerk. Gros-Schultze-Methode. Vergr. etwa 300fach

CEREBELLAR CORTEX: 400X ORIGINAL MAGNIFICATION, 15 μ SECTION, BIELSCHOWSKY SILVER TECHNIQUE



DR. JOSEPH J. WARNER

Climbing Fibers

Šplhavá vlákna

Olivo-cerebellar projections

(crossed)

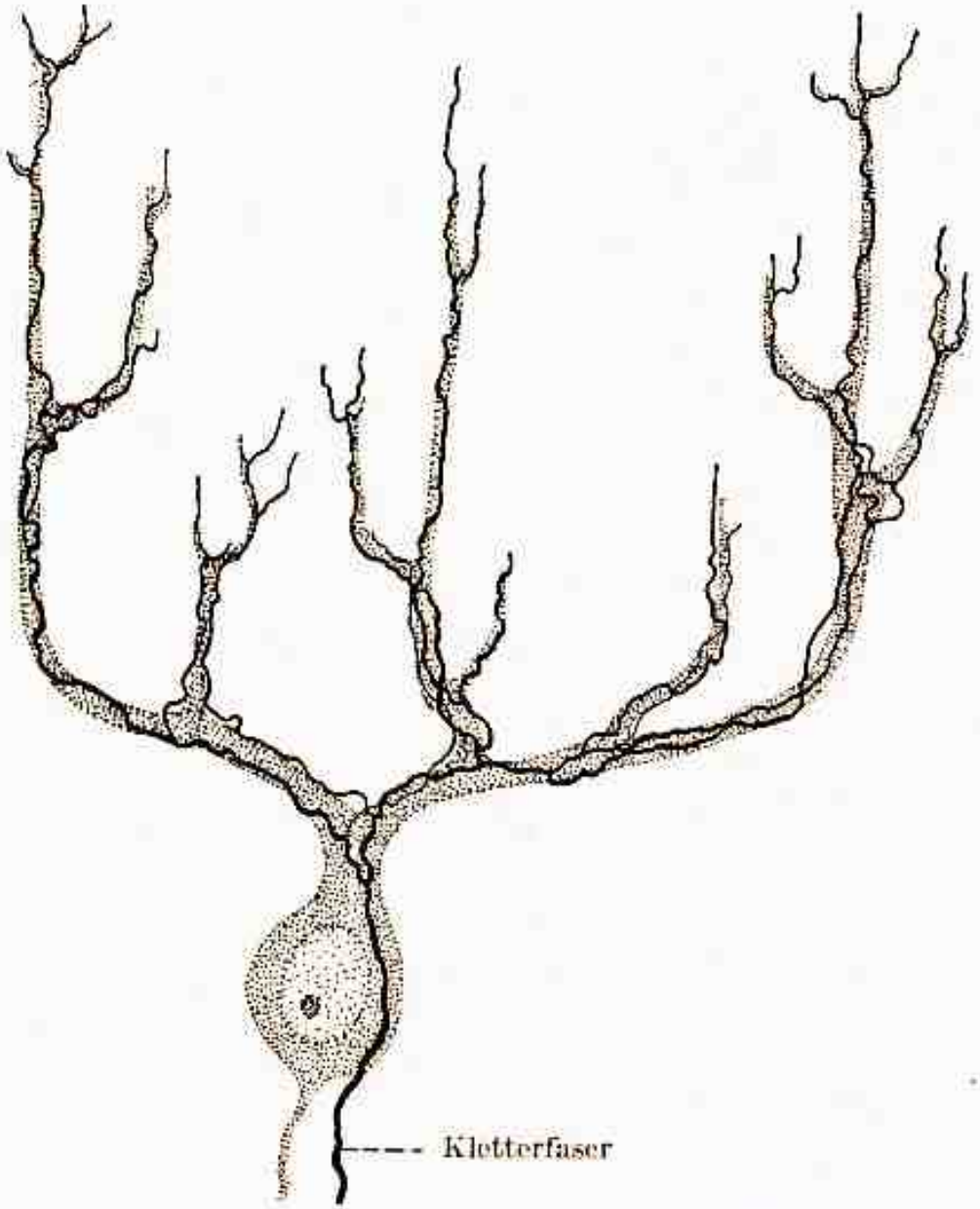
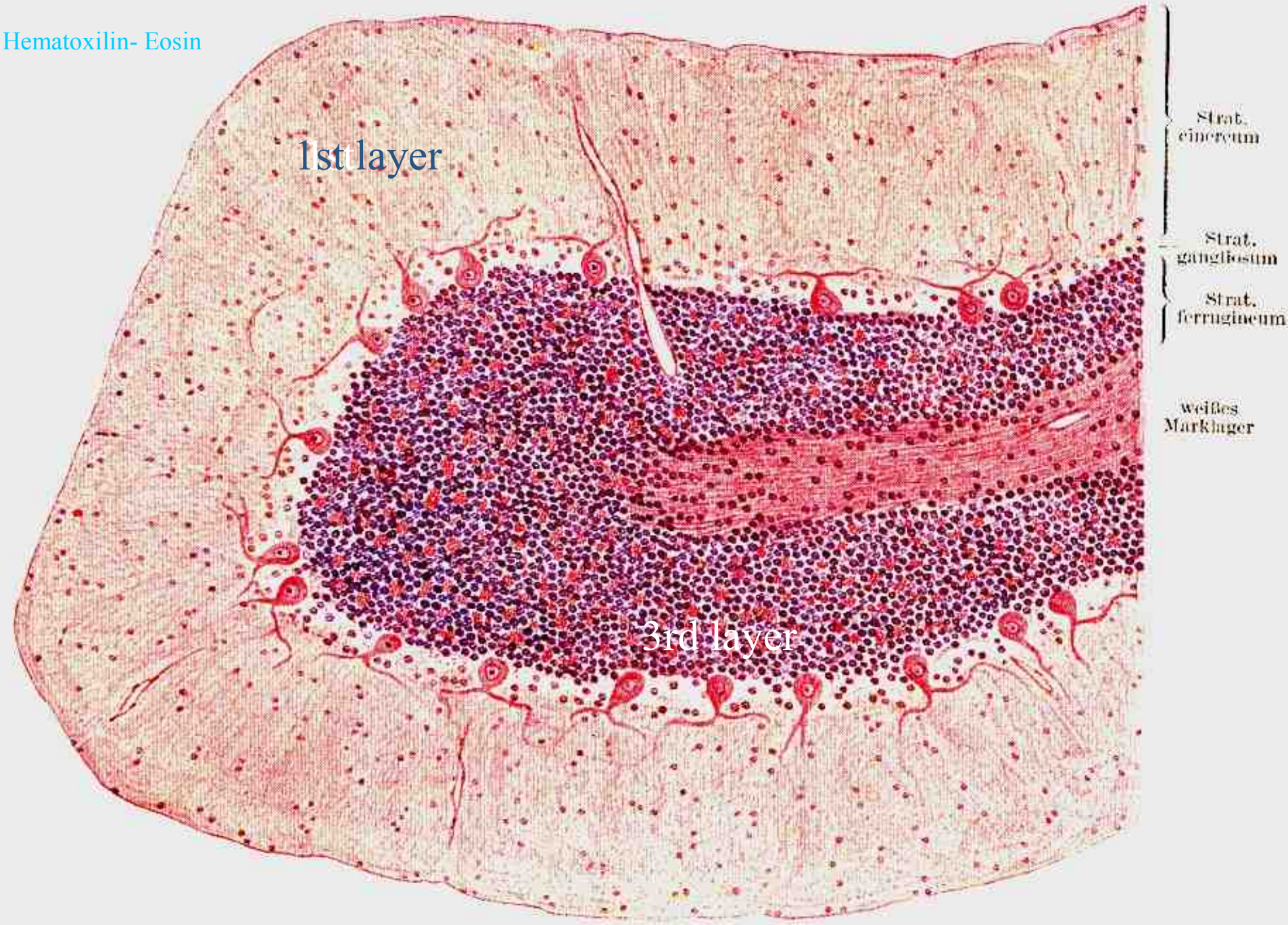


Abb. 264. Purkinjesche Zelle mit Kletterfaser (halbschematisch)

Hematoxilin- Eosin



Synaptophysine – immunostaining (pozitivita na synaptických kontaktech)

Image265. x4 synapto Švarc

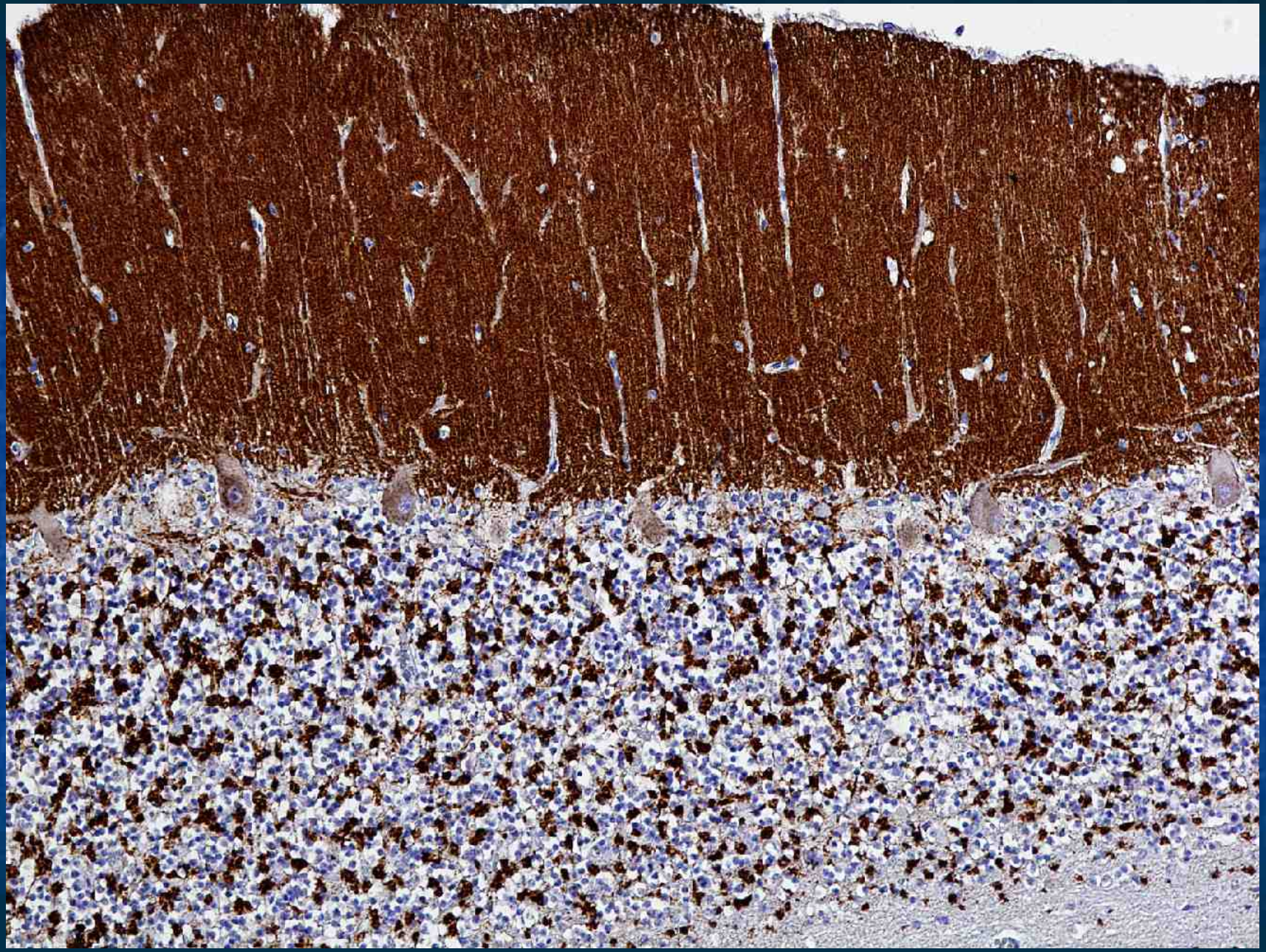


Image267.t4 x10

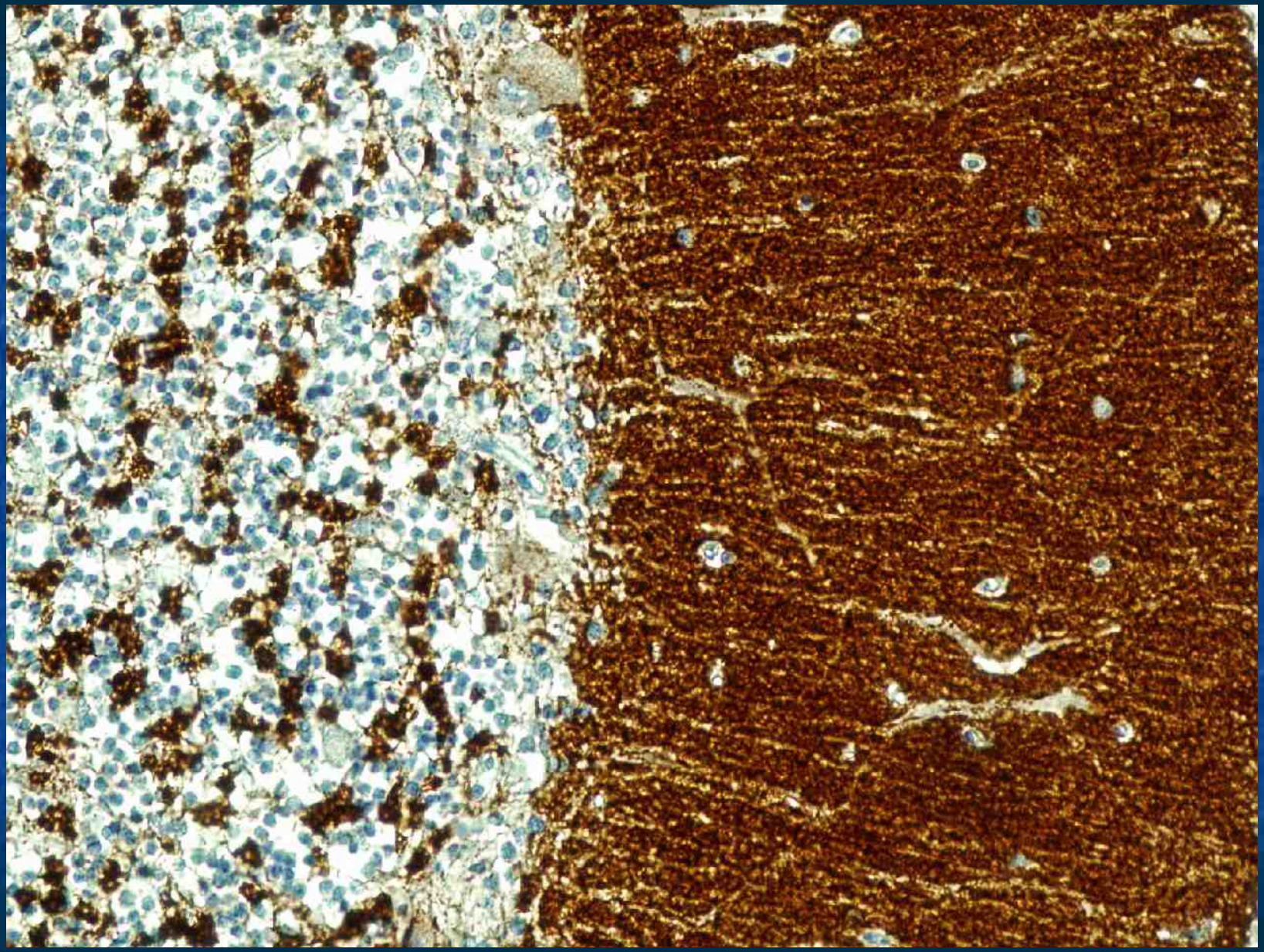
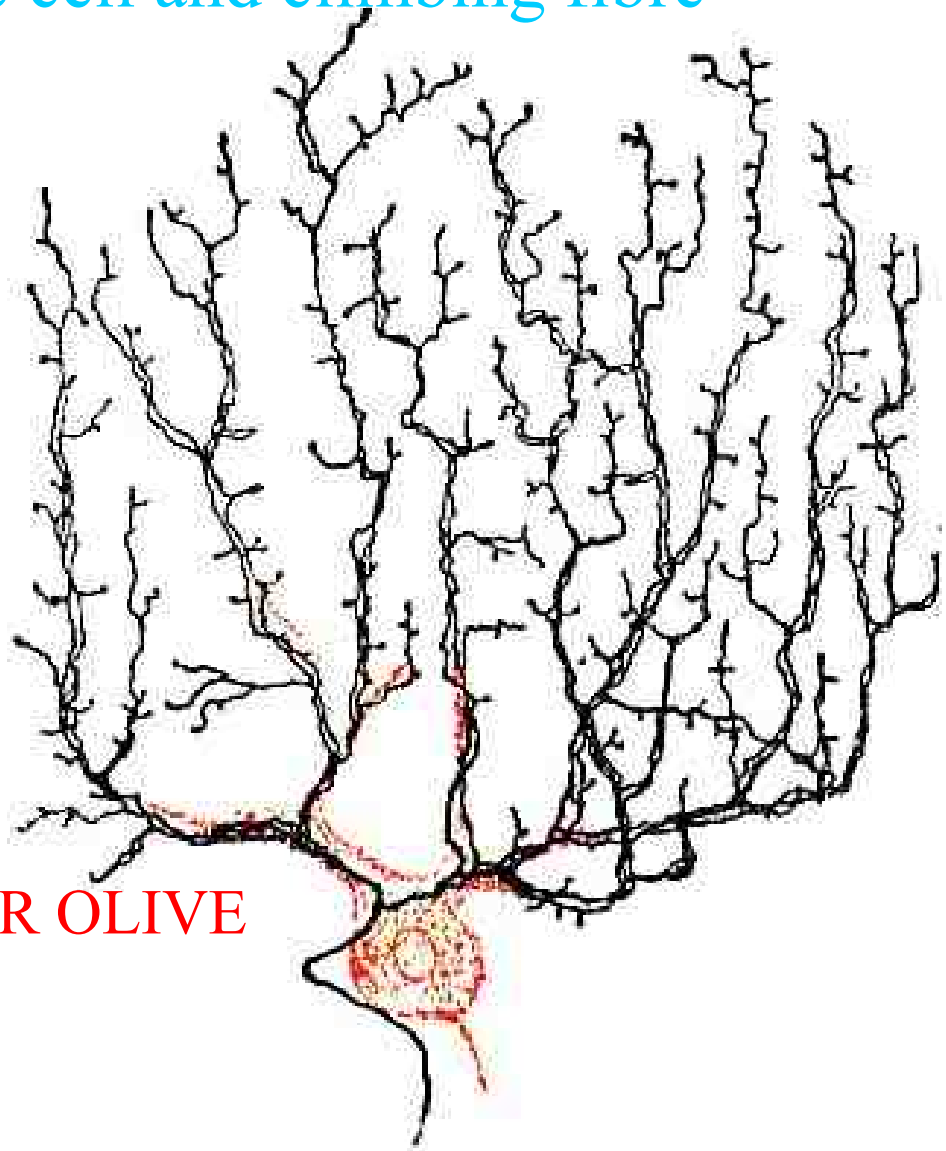


Image272 x20

Two types of afferent fibers
Dva typy aferentních vláken

- Šplhavá vlákna – inferior olive
- Mechová vlákna – spinal cord, brain stem (trigeminal ncc., vestibular ncc., RF, pontine ncc.)

Purkyně cell and climbing fibre



INFERIOR OLIVE

Terminals of the mossy fibers (cerebellar glomerulus – rosetta)

Zakončení mechového vlákna v granulární vrstvě (kontakt s dendrity několika granulárních buněk)

One mossy fiber has excitatory contacts with several granular cells

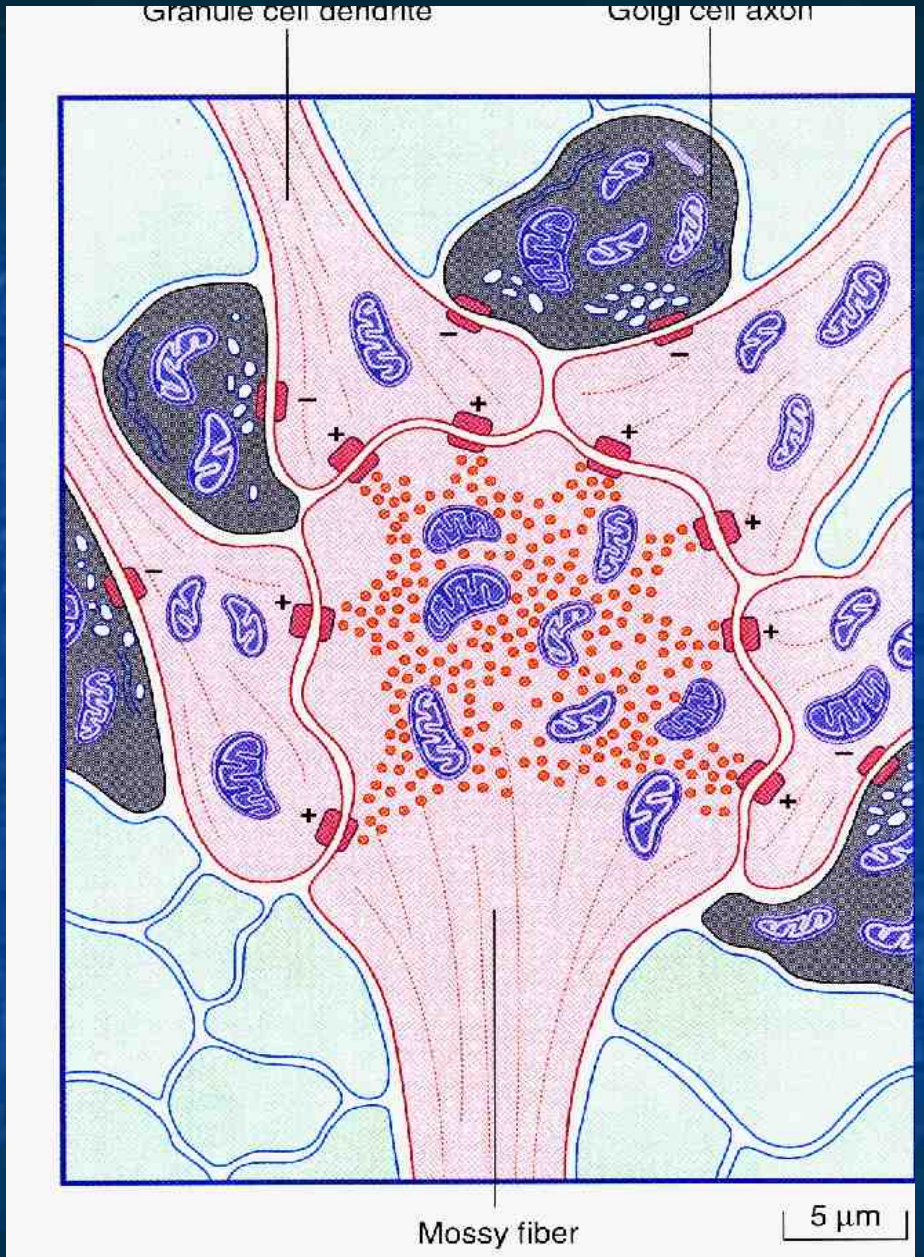
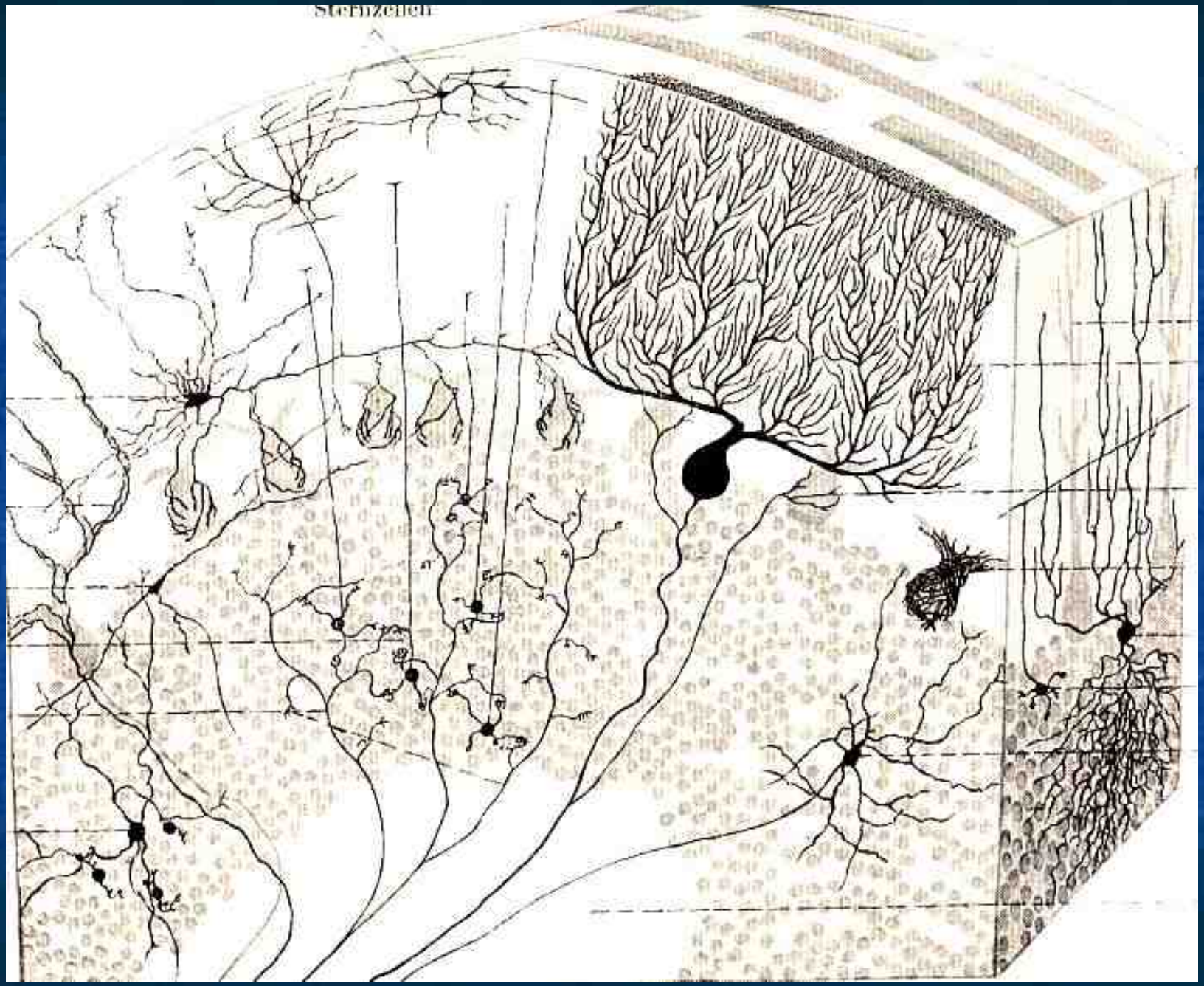


Figure 25.4 A synaptic glomerulus. +/– indicates excitation/inhibition.


Sternzellen

Mossy fiber

Mechová vlákna

↓

Granular cells

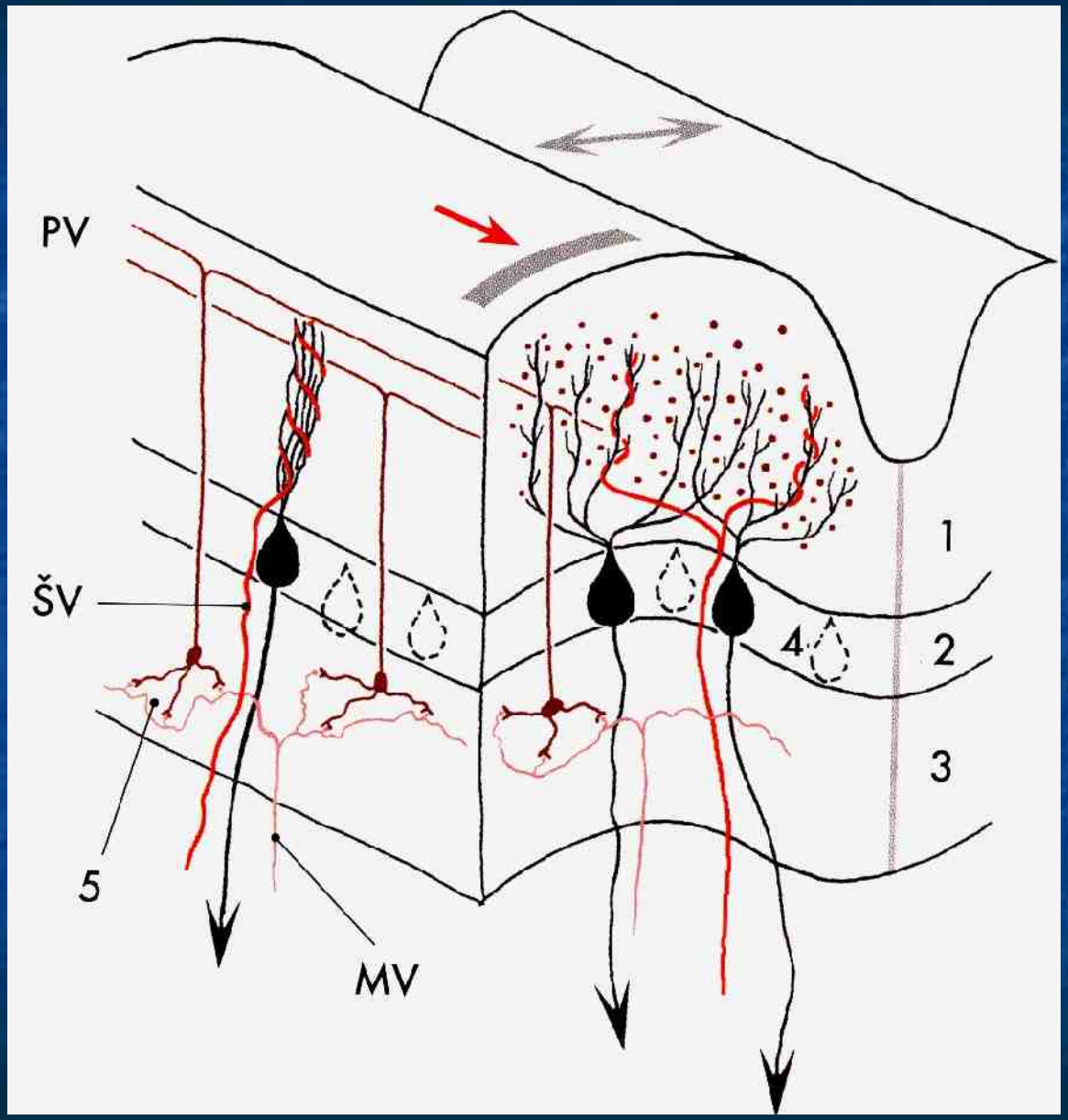
↓

Paralel fibers

↓

Purkyně cells
dendrites

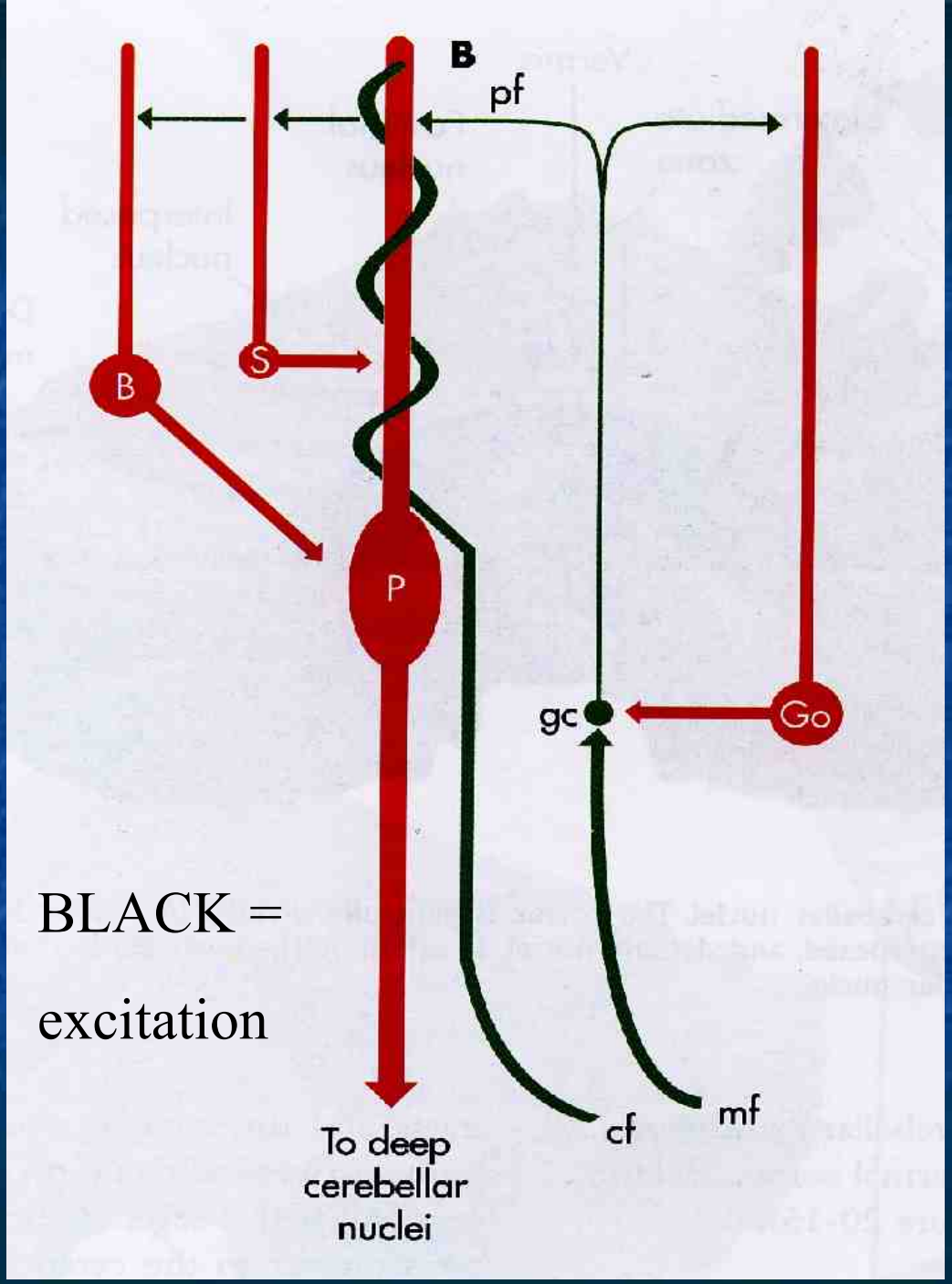
Excitatíon of P. cells =
inhibition of cerebellar
nuclei



RED = inhibition

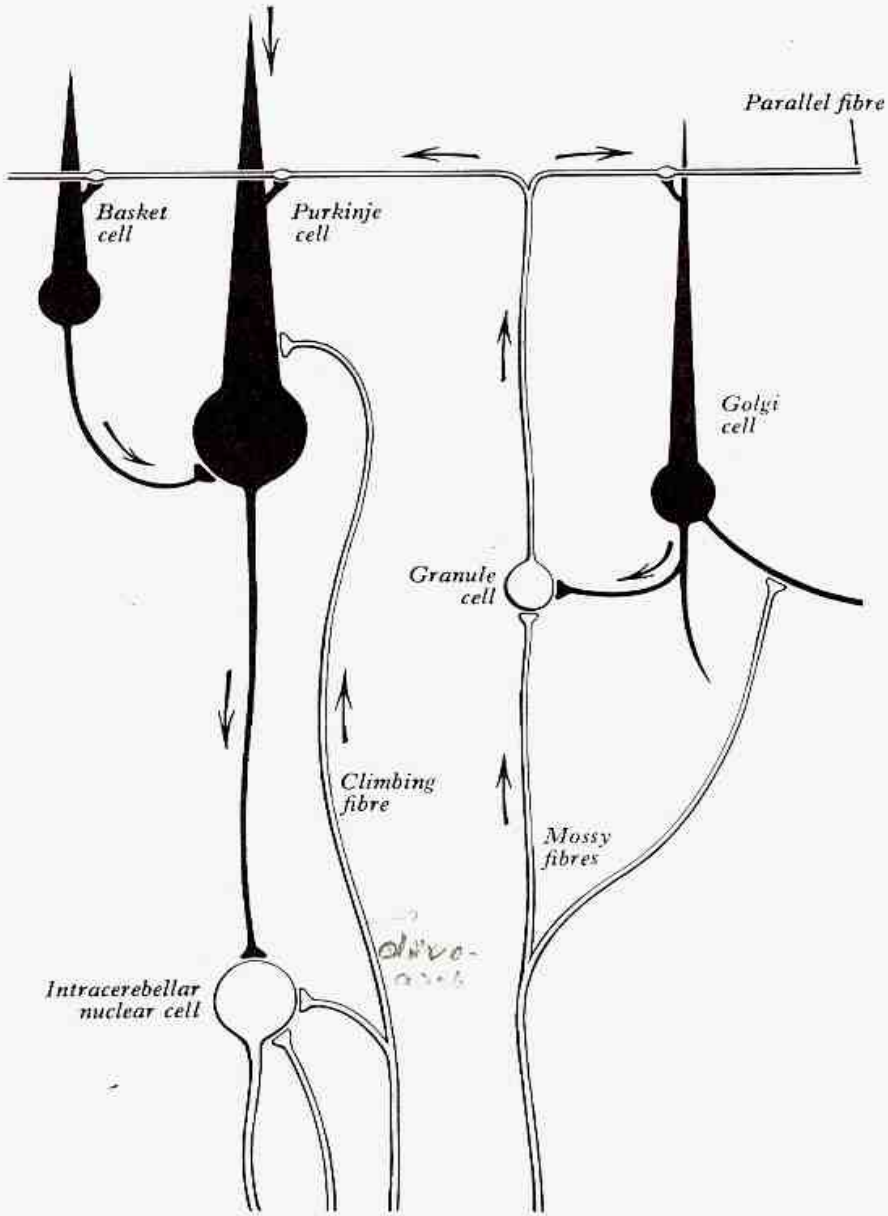
Červená = inhibice

Černá = excitace

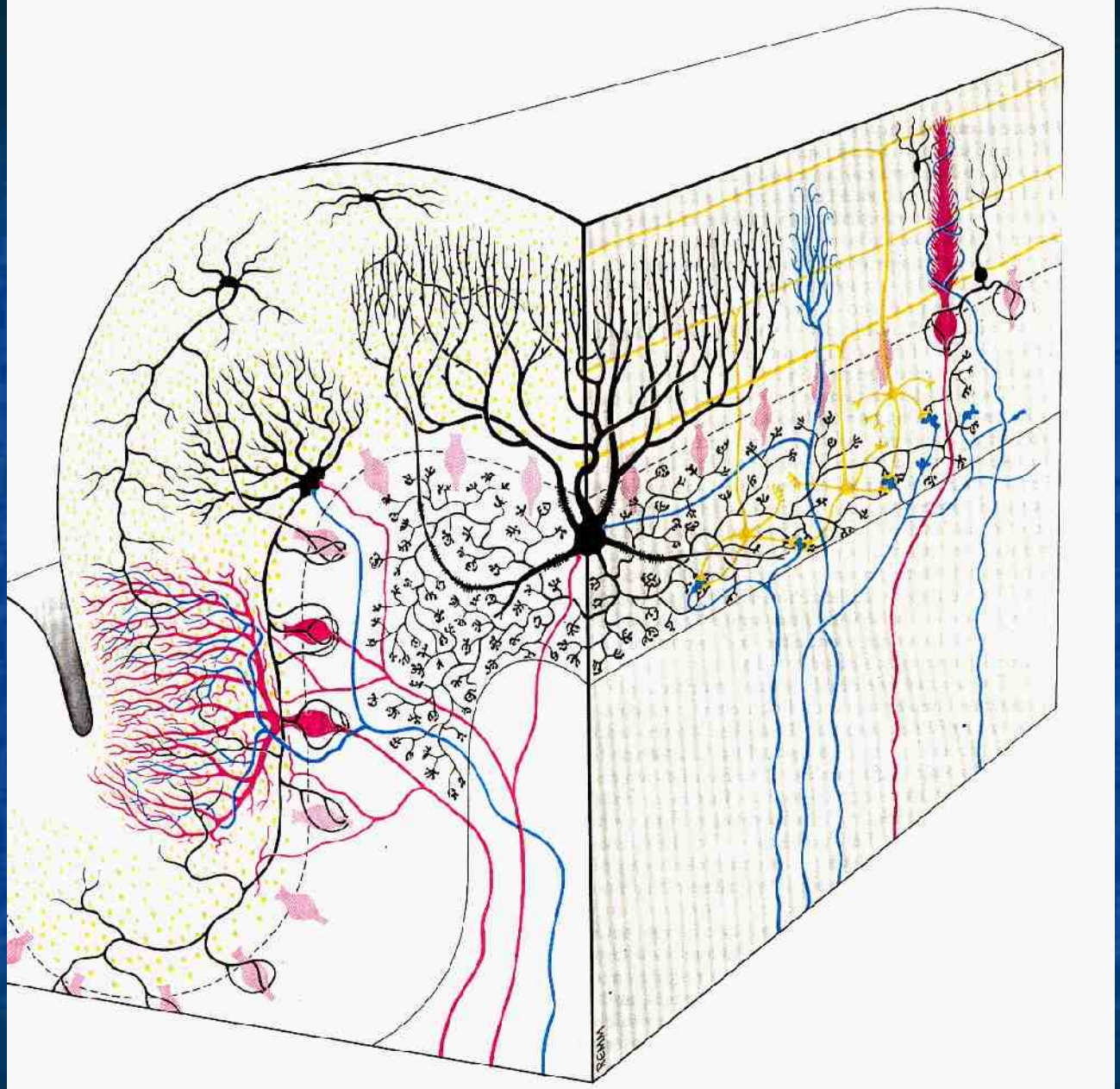


J. Eccles 1967

Nobel Prize in Physiology and medicine 1963



7.82 An analysis of the essential circuitry and synaptic contacts between the climbing and mossy afferent fibres, the main neuronal elements of the cerebellar cortex, and the neurons of the intracerebellar nuclei, based upon cytological and microelectrode studies. Excitatory cells, neurites and terminals are white surrounded by a black line; inhibitory elements are solid black. By courtesy of Professor J. C. Eccles.



79 The general organization of the cerebellar cortex: a single cere-
llar folium has been sectioned vertically, both in its longitudinal axis
ght part of the diagram) and transversely (on the left). Note: (1)
arkinje cells (red); (2) inhibitory interneurons (black) including outer
ellate, basket and Golgi cells; (3) granule cells and their ascending
ons which bifurcate into longitudinally disposed horizontal fibres
(yellow); (4) climbing fibres and mossy afferents (blue). Note also the
synaptic glomeruli formed between the terminals of the mossy afferent
fibres, the complex dendrite tips of the granule cells, and the ramifica-
tions of the Golgi cell axon. (Redrawn from: *The Cerebellum as a Neuronal Machine* by J. C. Eccles, M. Ito and J. Szentágothai. With the permission of the authors and the publishers Springer, 1967.)

Mossy fibers –

Granule cells-

Paralel fibers –Purkyně cells

Climbing fiber –

Purkyně cell

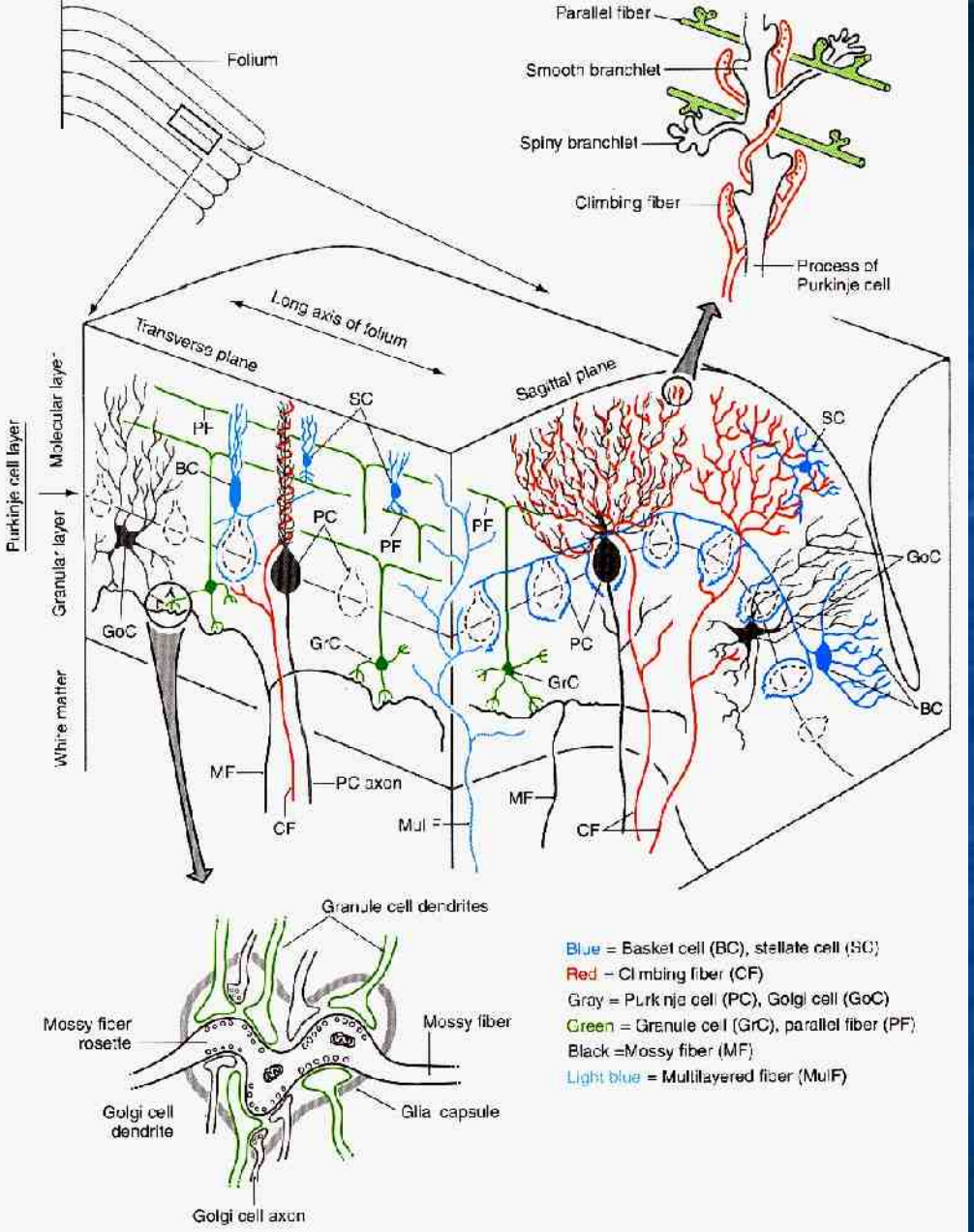


Figure 27–8. Cell types and synaptic relations in the cerebellar cortex in transverse and sagittal planes. Note the struct the cerebellar glomerulus (*lower left*) and the interaction of parallel and climbing fibers (*upper right*) with the dendritic pro of Purkinje cells. Compare with Figure 27–9.

AFFERENT AND EFFERENT CONNECTIONS

- Tracts : **Mossy fibers**
- Vestibulocerebellar (from the labyrinth and vestibular nuclei)
- Spinocerebellar ant., post.,
- rostral, cuneocerebellar
- Reticulocerebellar
- Nucleocerebellar
- Pontocerebelar !! (cortico-ponto-cerebellar)
- **Climbing fibers**
- **Olivocerebellar fibers**

- Corticonuclear (from the cerebellar cortex to the nuclei)
- Vermis – nc. fastigii
- Paravermal zone – nc. embol. Nc. glob.
- Lateral hemisphere – nc. dentatus

- From cerebellar nuclei to the brain stem and to the thalamus

Spinocerebellar Pathways

Posterior T1 – L2, uncrossed

ICP, proprioceptors

Anterior L3 – L5

Crossed, SCP, cutaneous signals

Rostral C4 – C8

Uncrossed, ICP

Cutaneous signals

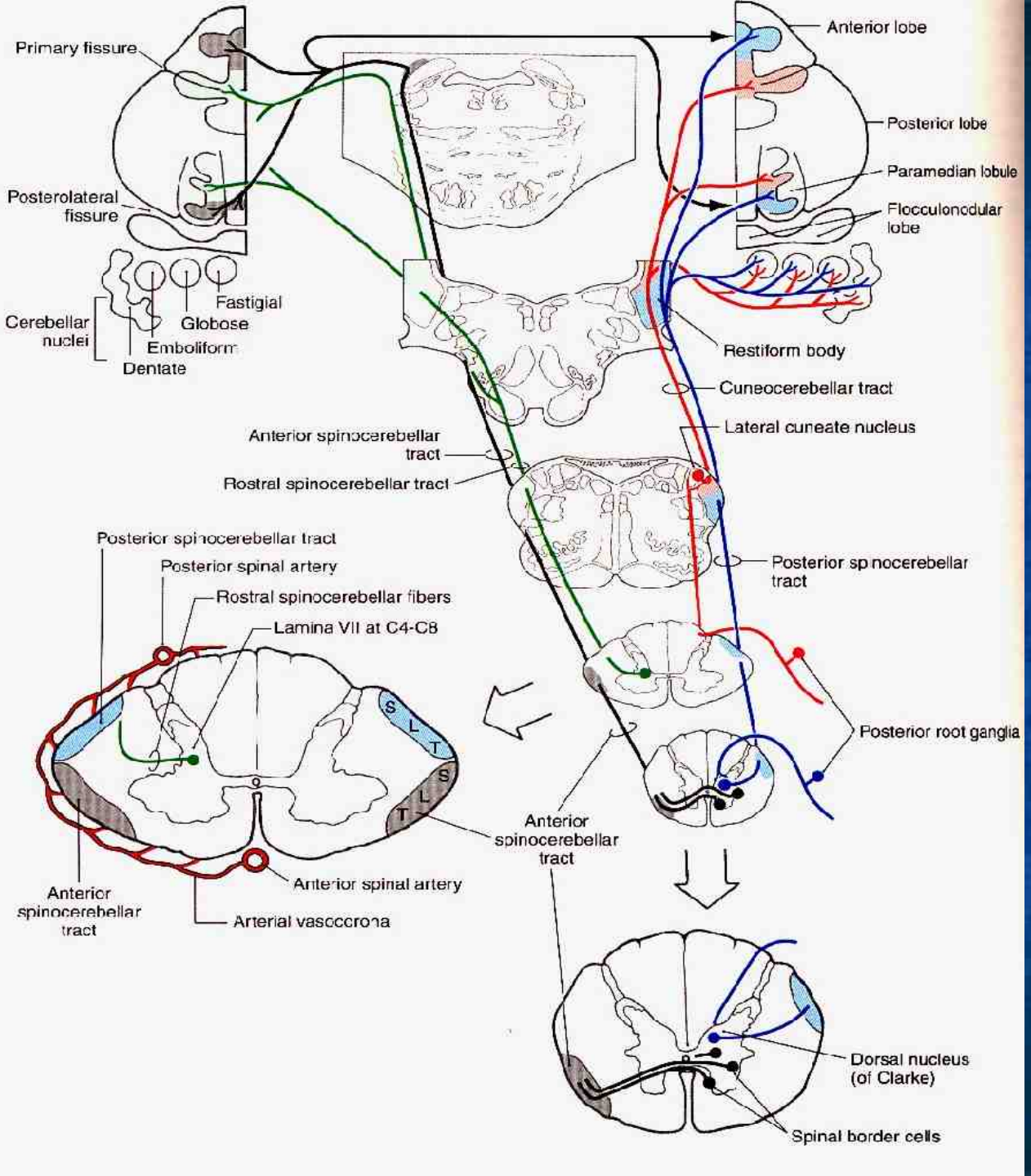
Cuneocerebellar

C2 – T4

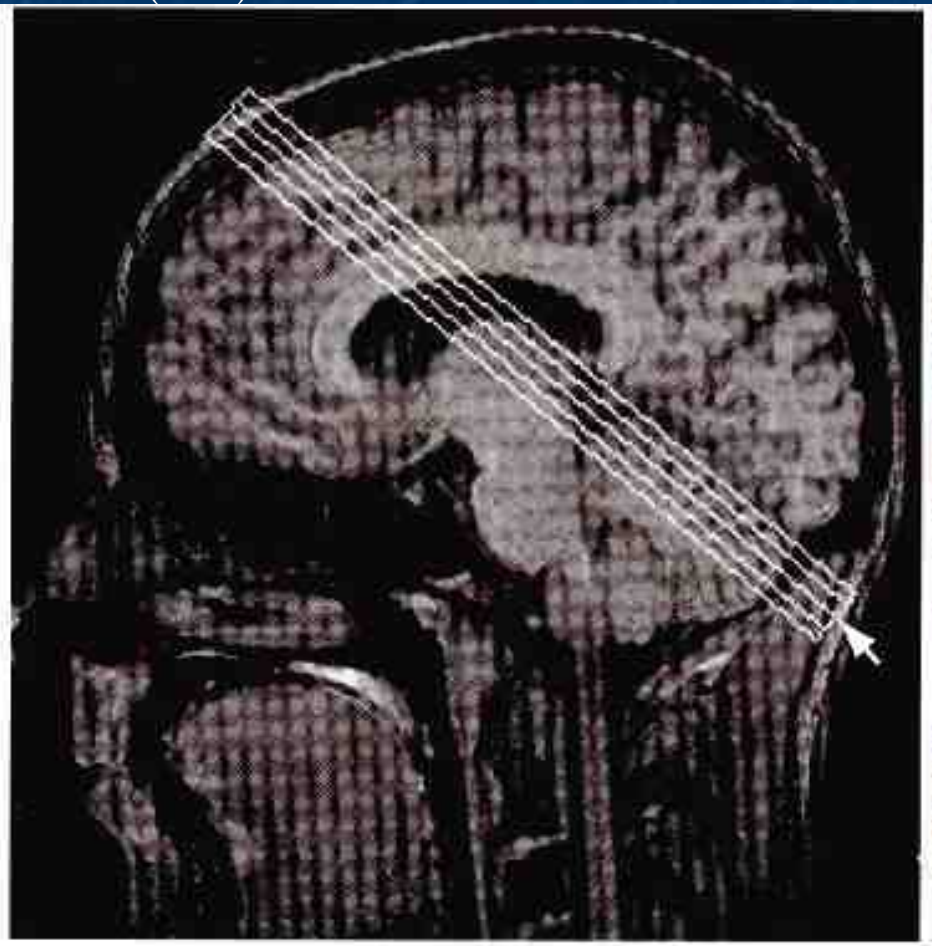
uncrossed

Lateral cuneate nc., propriocept.

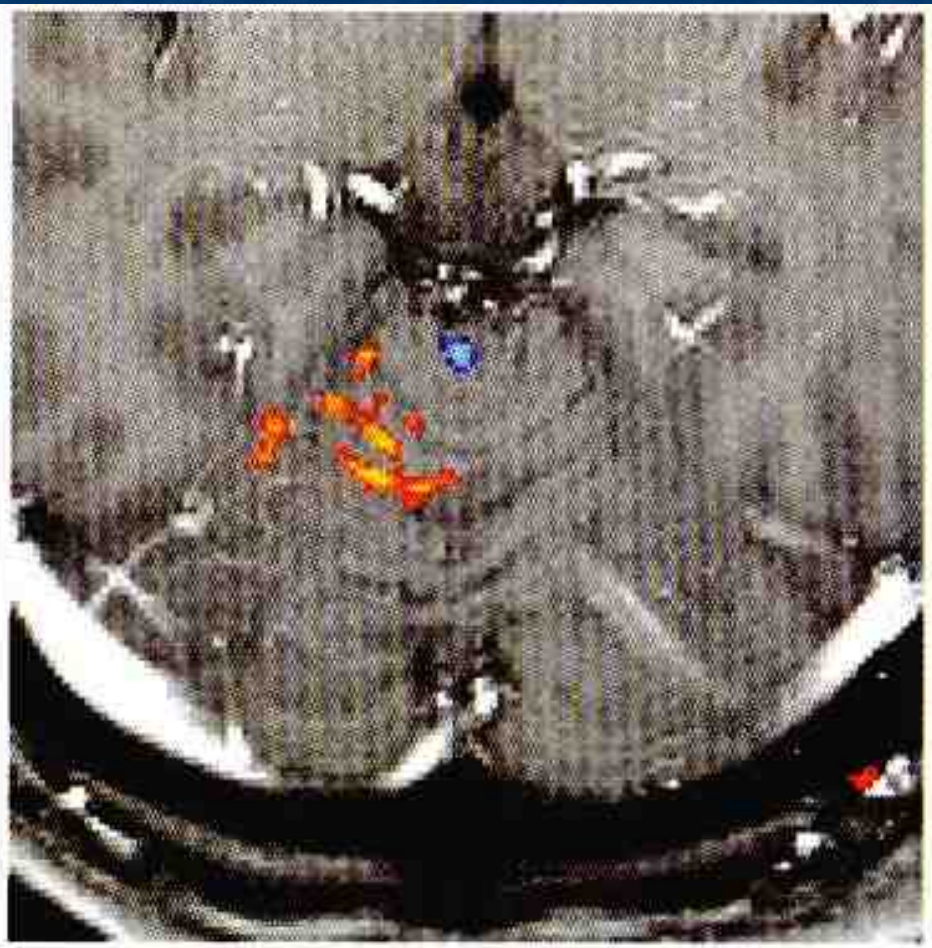
ICP



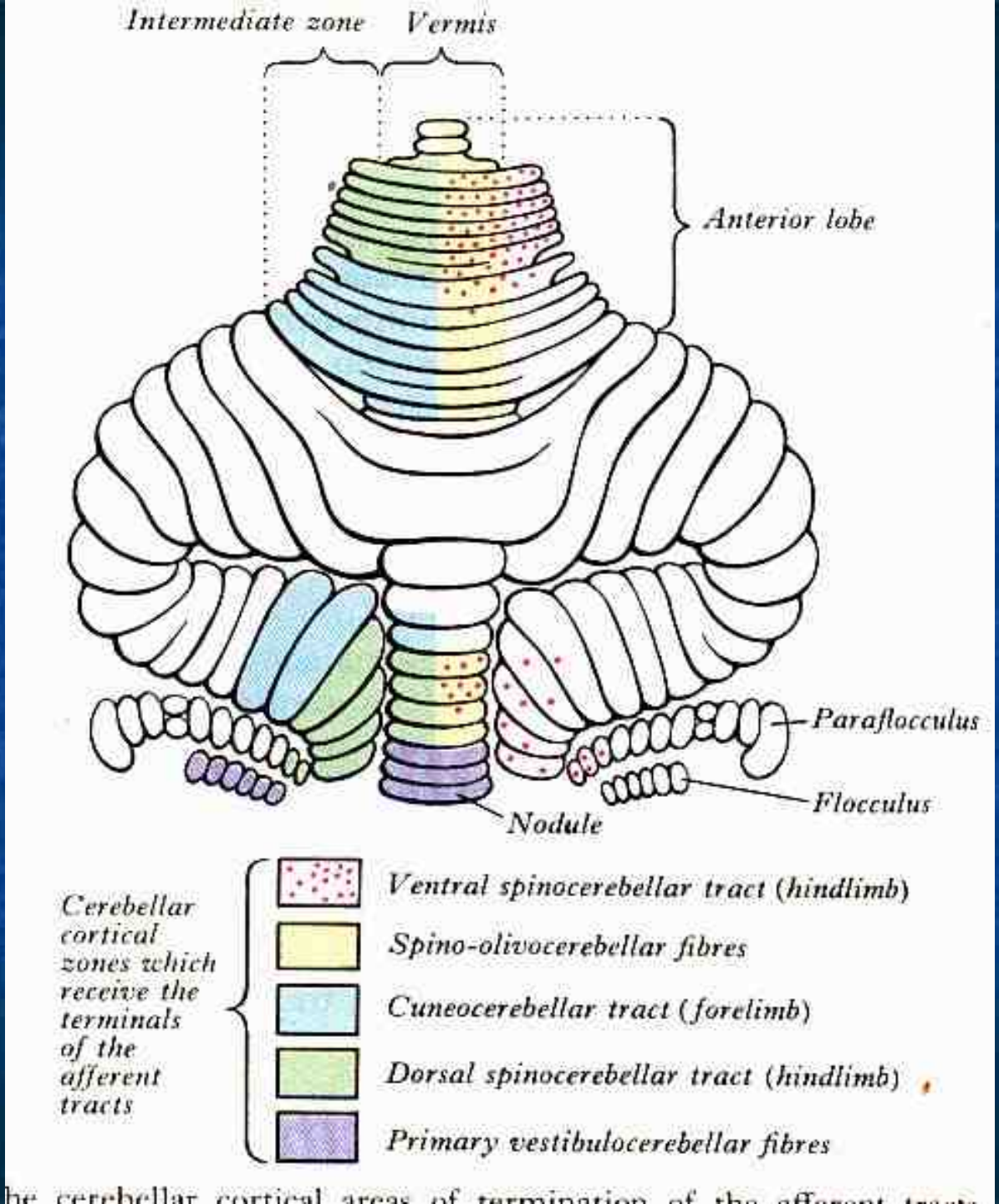
FMI – areas of increased blood flow during flexion and extension of ipsilateral hand (red, orange) and foot (blue)



Somatotopy in the cerebellar cortex



Afferent projections



he cerebellar cortical areas of termination of the afferent tracts

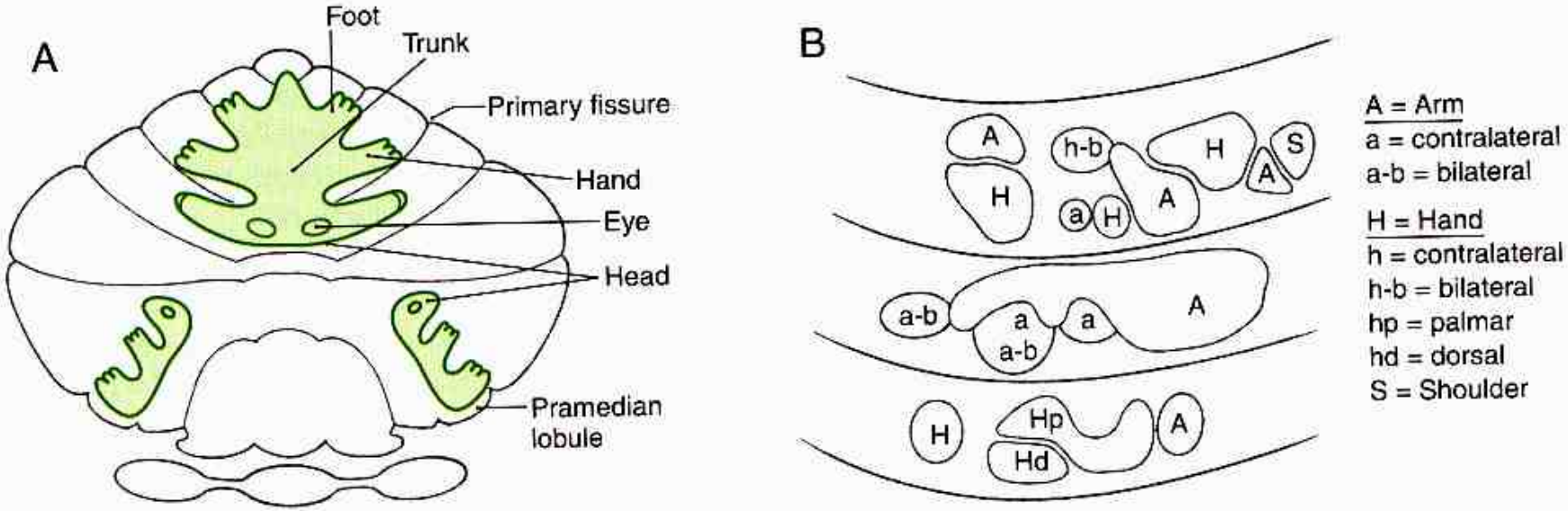
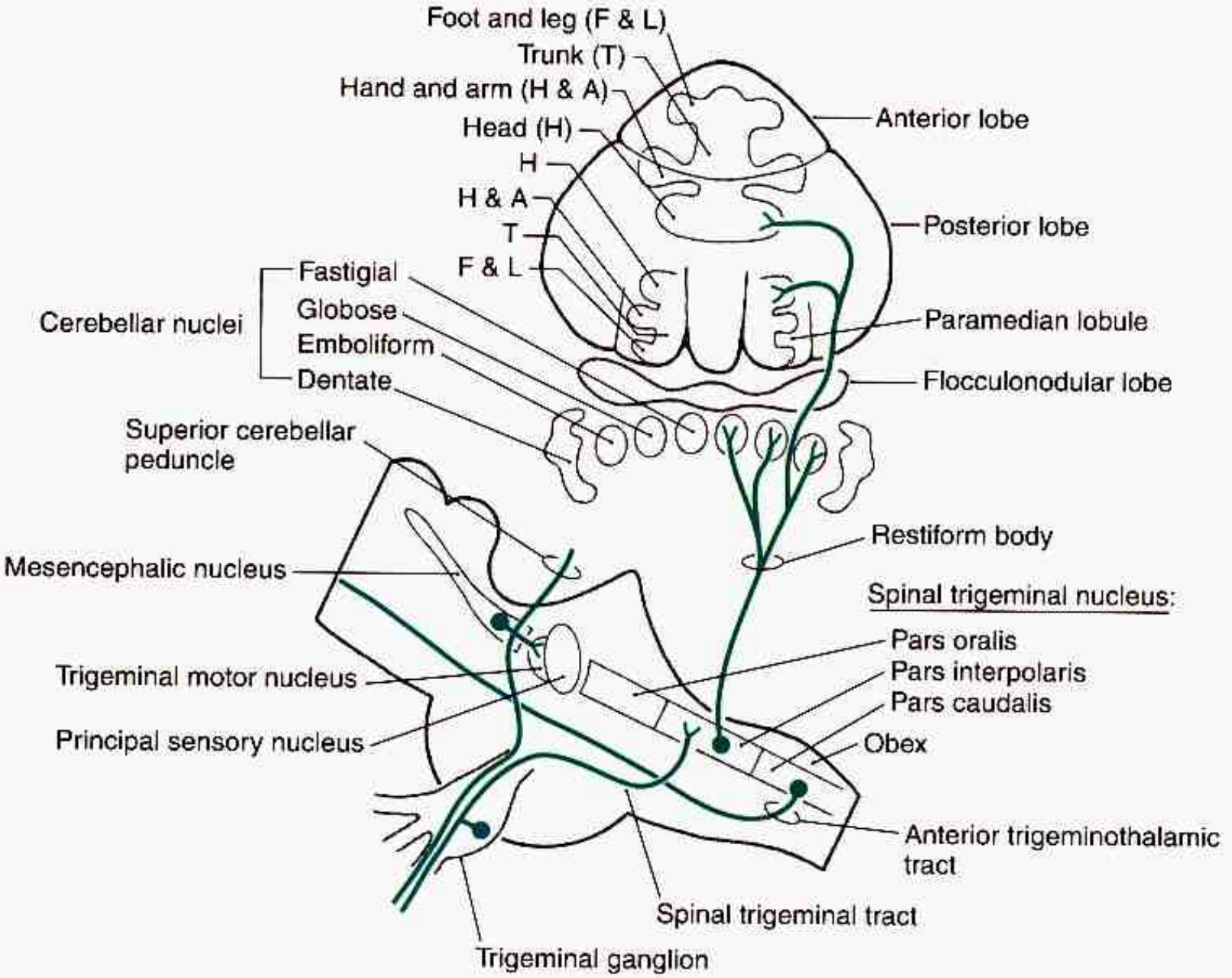


Figure 27–10. Somatotopy in the cerebellar cortex (*A*) and a summary representation of fractured somatotopy in the paramedian lobule (*B*) of a primate. In the somatotopic map, body areas were originally thought to be continuous (*A*), but more recent studies suggest that discontinuous body parts (or areas) may be represented in immediately adjacent cortical regions (*B*). (*B* is adapted from Welker et al, 1988, with permission.)

Trigeminocerebellar Connections



Cortico- pontine pathway -

17 millions fibers

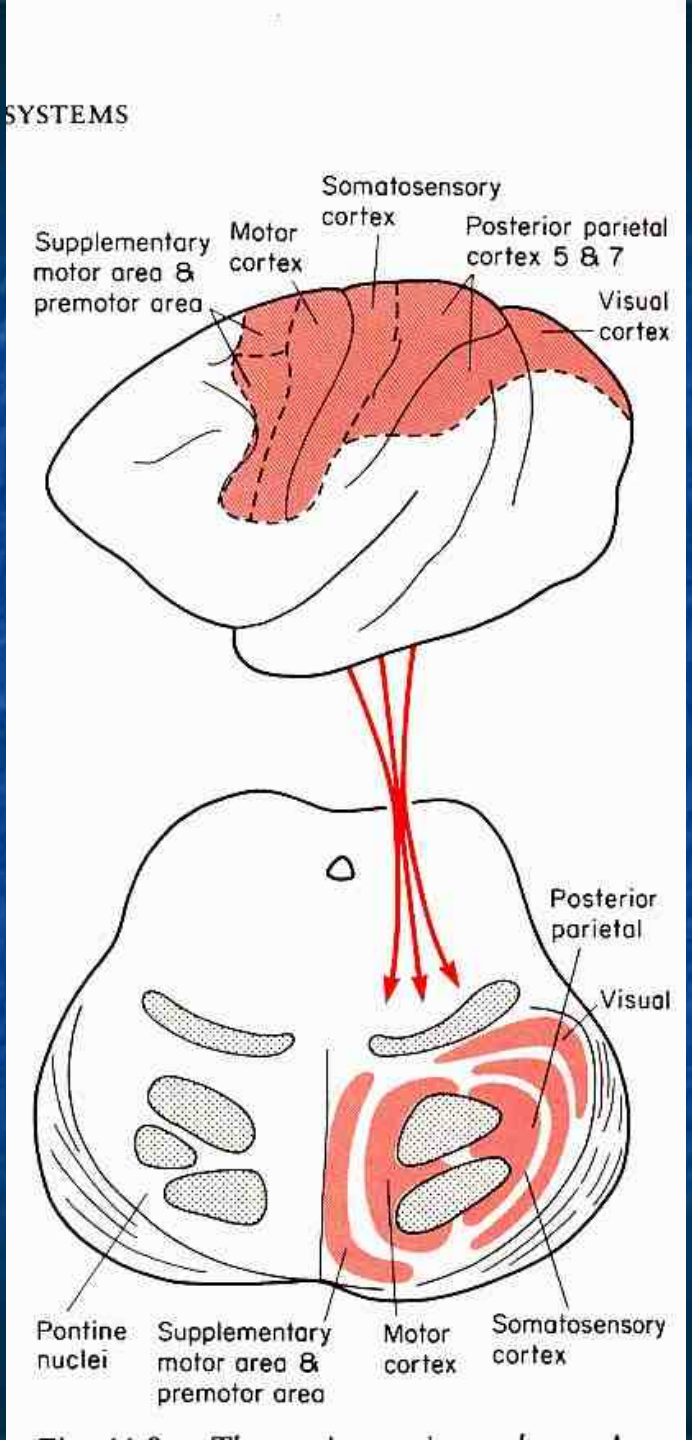
Neocortex –

ipsilat. pontine ncc. - pontocerebellar pathway –

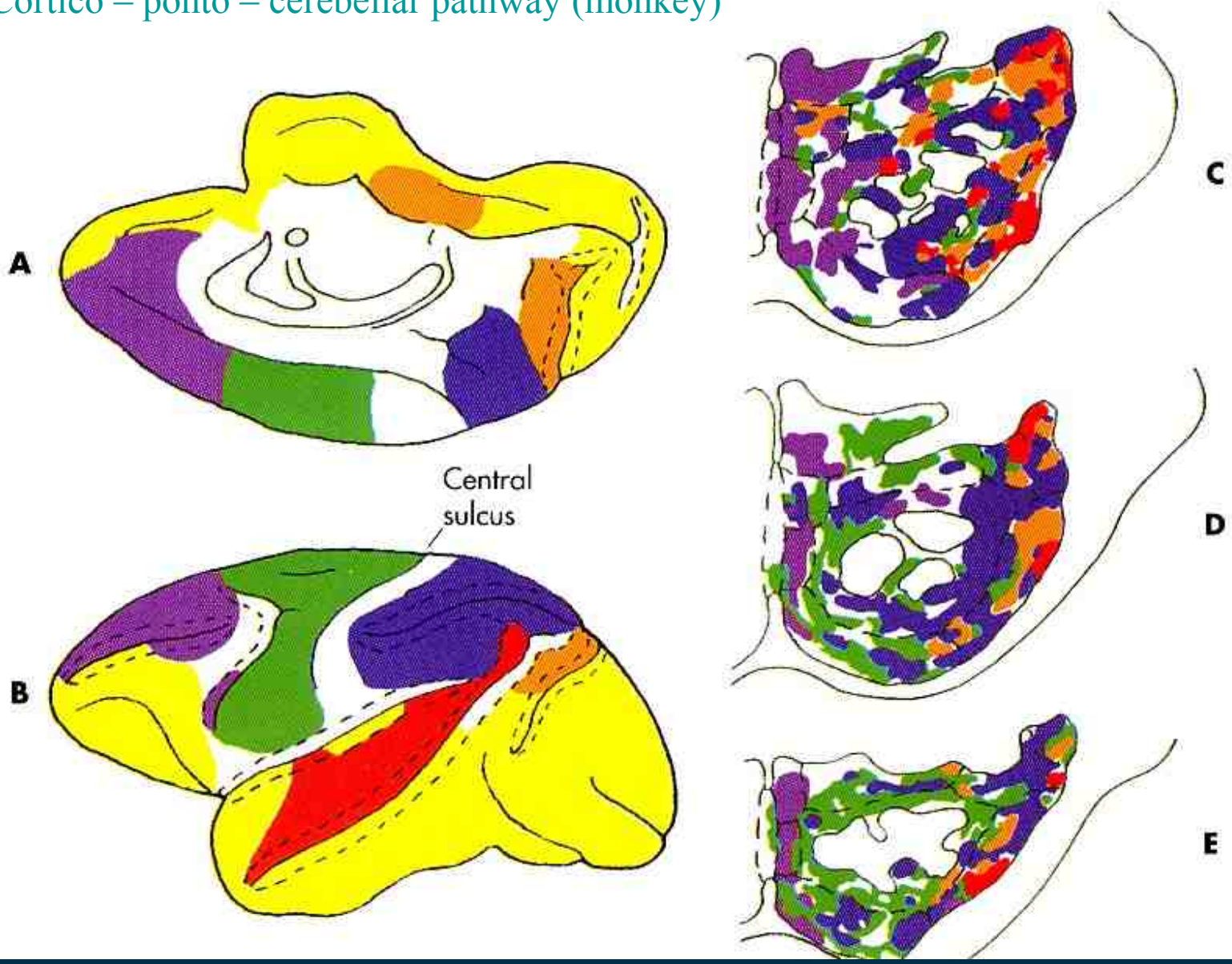
contralateral cerebellar cortex (mossy fibers)

Cortex – ncc. Pontis –

Mozečková kůra

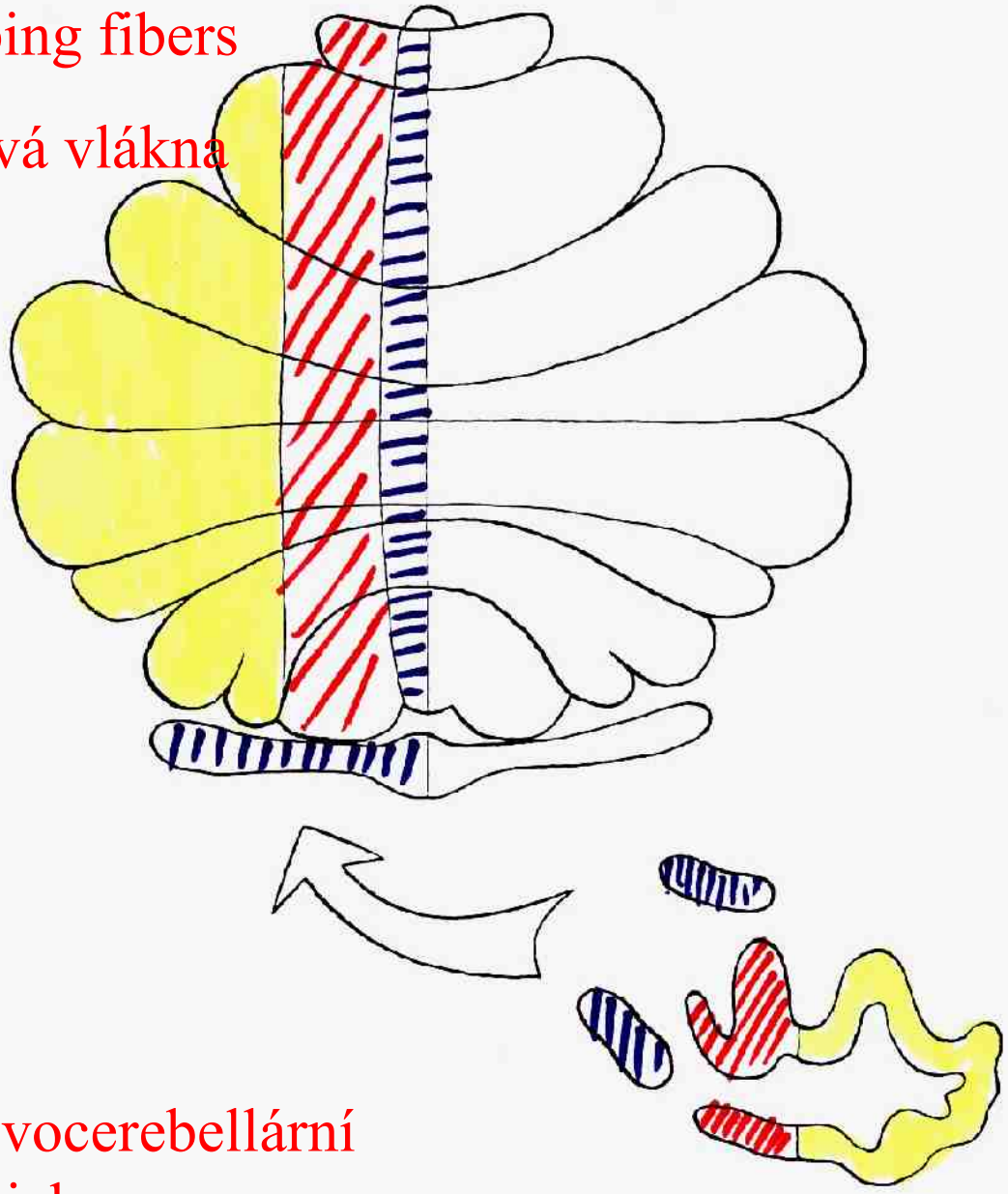


Cortico – ponto – cerebellar pathway (monkey)



Climbing fibers

Šplhavá vlákna



Olivocerebellární projekce

Inferior olive

AFFERENTS TO THE CEREBELLAR CORTEX I.

- **Climbing fibers** – inferior olive (each P.cell receives only 1 c.f., many synapses with P.c.), excitatory (glutamate), firing frequency of the c.f. is very low (1 impulse/sec), c.f. elicit burst of action potentials in the P.c.
- C.f. inform about errors in the execution of movements – error indicators !!

AFFERENTS TO THE CEREBELLAR CORTEX II

- **Mossy fibers** - spinal cord, RF, pontine nuclei, ncc. cranial nerves.
- End in the granular layer and each of which contacts large number of granular neurons. Granular cell axon contacts large number of P. c. via paralel fibers.
- Mossy fibers are excitatory (glutamate).
- Each mossy fiber influences many P.c. but the excitatory effect is weak. Many mossy fibers must be active together to provide sufficient excitation to fire a P.c.
- Mossy fibers provide precisely graded information about movements, skin stimulations, joint position and about motor comands issued from the cerebral cortex.

EFFERENT CONNECTIONS OF THE CEREBELLAR CORTEX

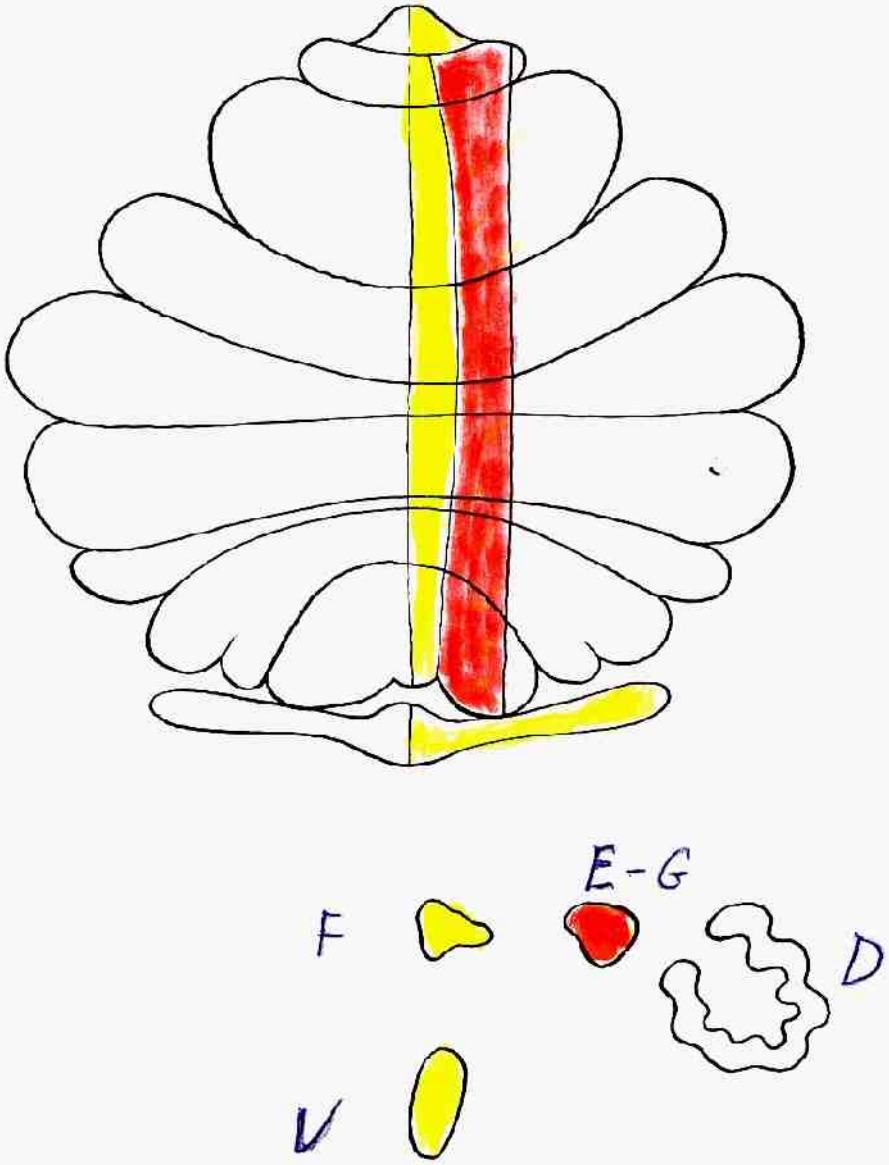
CEREBELLAR CORTEX – CEREBELLAR NUCLEI

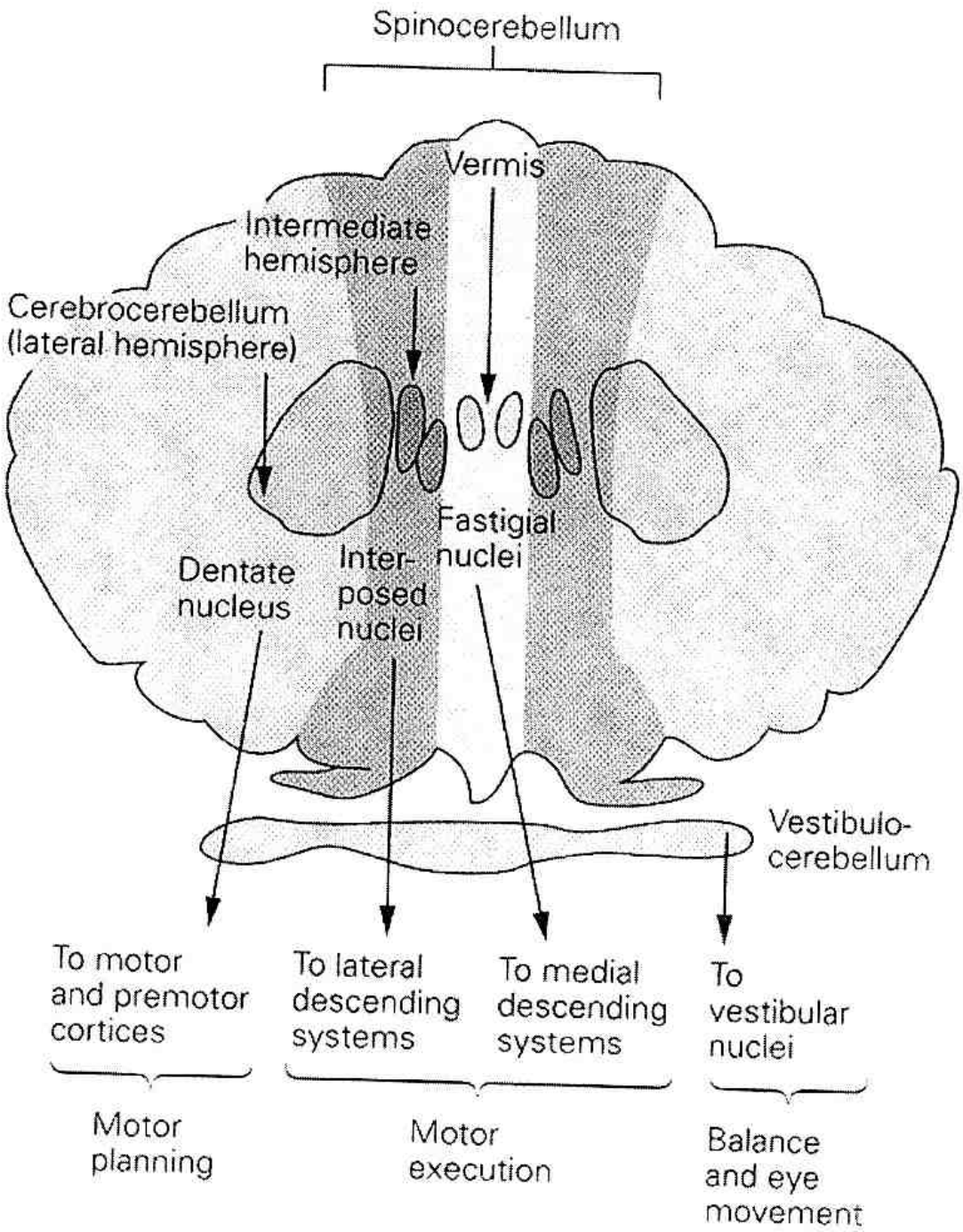
VERMIS – NC. FASTIGII, VESTIBULAR NCC.

PAVERMAL ZONE – EMBOLIFORMIS, GLOBOSE NCC.

LATERAL HEMISPHERE – DENTATE NC.

Kortikonukleární projekce




Spinocerebellum
Vermis
Intermediate hemisphere
Cerebrocerebellum (lateral hemisphere)
Fastigial nuclei
Dentate nucleus
Inter-posed nuclei
Vestibulo-cerebellum
To motor and premotor cortices
To lateral descending systems
To medial descending systems
To vestibular nuclei
Motor planning
Motor execution
Balance and eye movement

EFFERENT CONNECTIONS OF THE CEREBELLAR NUCLEI

- **Fastigial nucleus** – vestibular nuclei, reticular formation
- **Emboliformis + globosus nucleus** -.reticular formation, nc. ruber, thalamus
- **Nucleus dentatus –** nc. ruber, contralateral thalamus (**ventrolateral nucleus,** intralaminar thalamic nuclei, ventral anterior nc.,

- **Ventrolateral nucleus – primary motor cortex (area 4)**

Dentato-thalamic projection

(V. Chan-Palay)

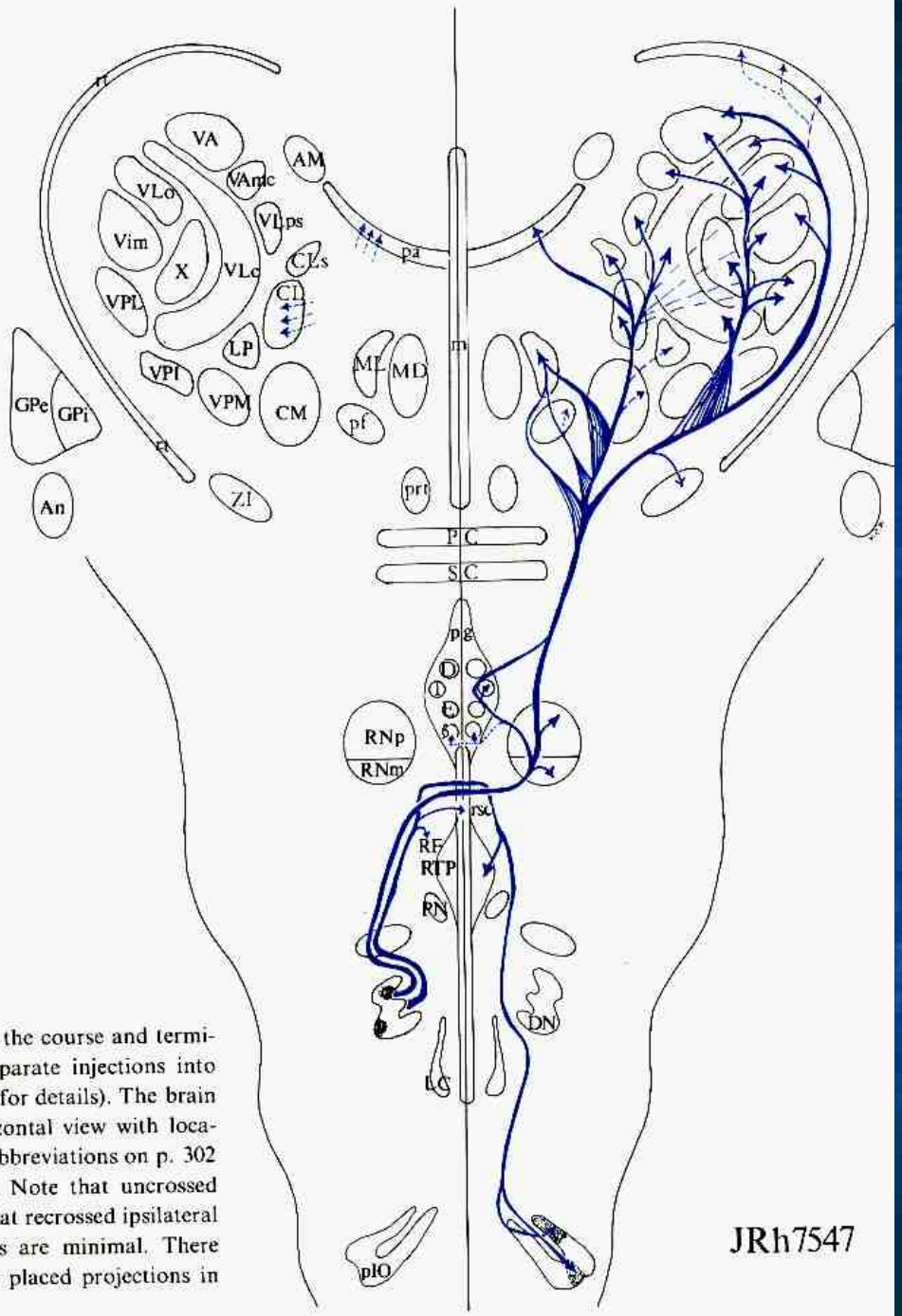


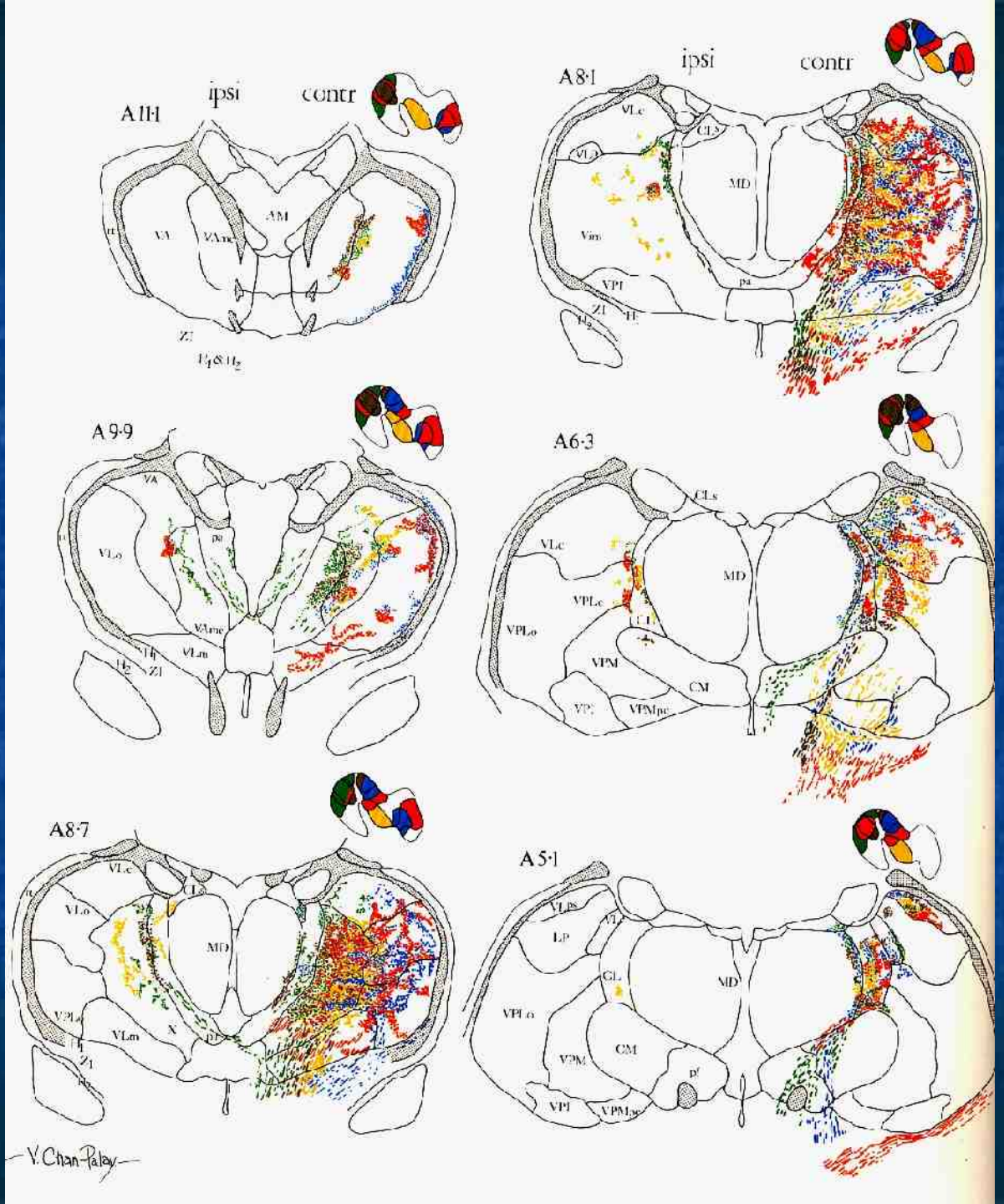
Fig. 13-7c. Summary diagram showing the course and terminations of labeled axons after two separate injections into the dentate nucleus, JRh 7547 (see text for details). The brain stem and thalamus are shown in horizontal view with locations of nuclei indicated according to abbreviations on p. 302 (Compare with Figs. 13-4c to 13-8c). Note that uncrossed ipsilateral projections are absent and that recrossed ipsilateral projections to the ipsilateral thalamus are minimal. There is a pronounced emphasis on laterally placed projections in the thalamus

Thalamus

Nc. Ventralis Lateralis

Nc. VL – Motor cortex

V. Chan-Palay



V. Chan-Palay :

Cerebellar cortex

Dentate nucleus

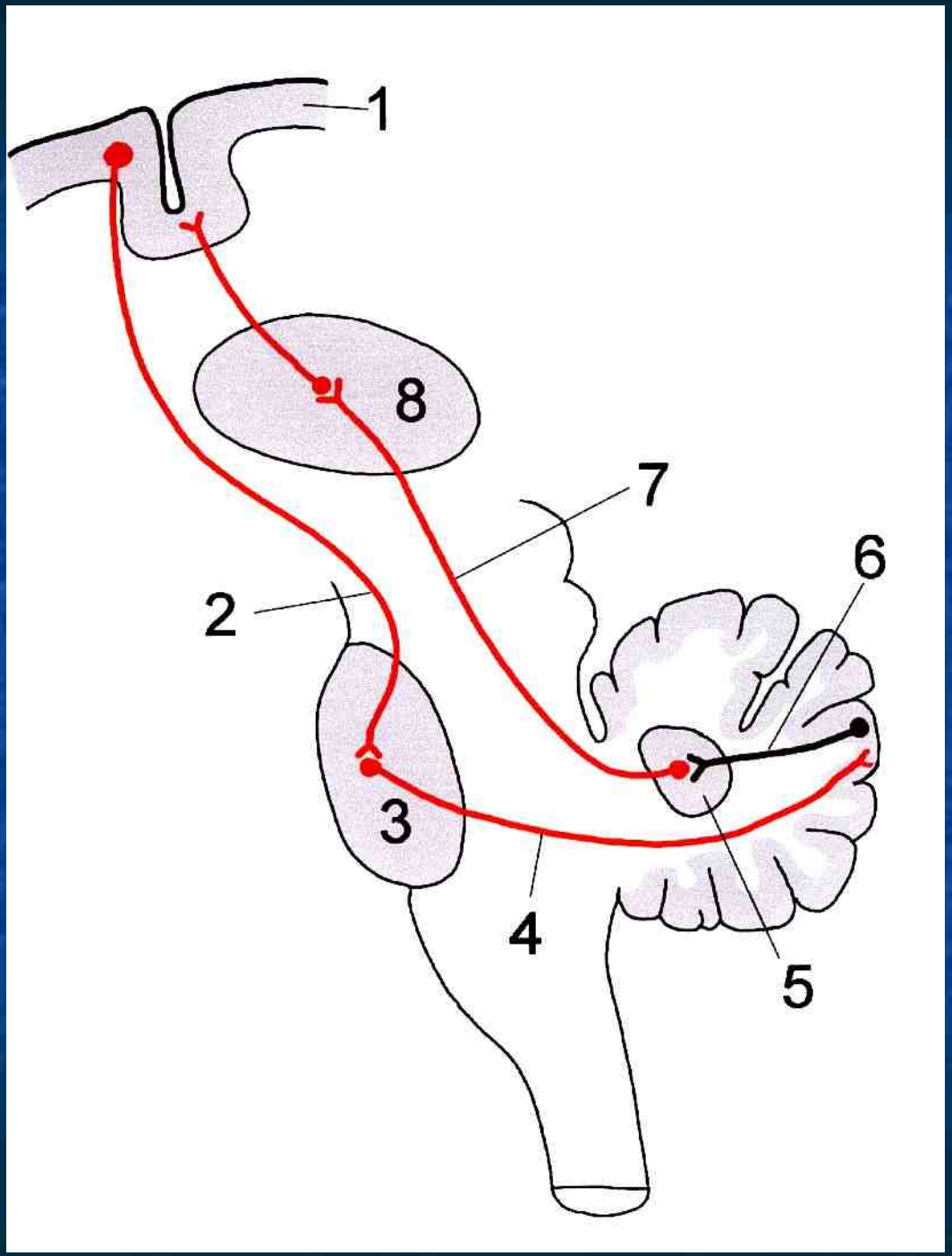
Victoria Chan-Palay, author and illustrator, attended Smith College and holds a Ph.D. in Anatomy from Tufts University School of Medicine. The present volume is an expanded version of an honours thesis which was presented to the Harvard Medical School at the conclusion of her clinical studies in 1975. It earned for her the degree of M.D. *summa cum laude*, a rare distinction at that institution. She also received the Leon Reznick Memorial Prize for excellence and accomplishment in research at the Harvard Medical School the same year.

Dr. Chan-Palay is a native of Singapore, a champion swimmer, an ardent supporter of equal rights for all women, the author of numerous articles in neuroscience journals and the co-author and illustrator of the book "Cerebellar Cortex – Cytology and Organization". This book was one of the "Fifty Best Books in 1974 in the Federal Republic of Germany, judged according to typography, printing, illustration and binding". It was awarded a special citation for its jacket design and received the 1975 Award for the Best Book for Professional Readership given by the American Medical Writers Association.

Cortico-ponto-cerebello-cortical circuit

(Transcerebellar Circuit)

3 x



1 –neocortex

3- pontine ncc.

5 – cerebellar ncc.

8 – thalamus (VL)

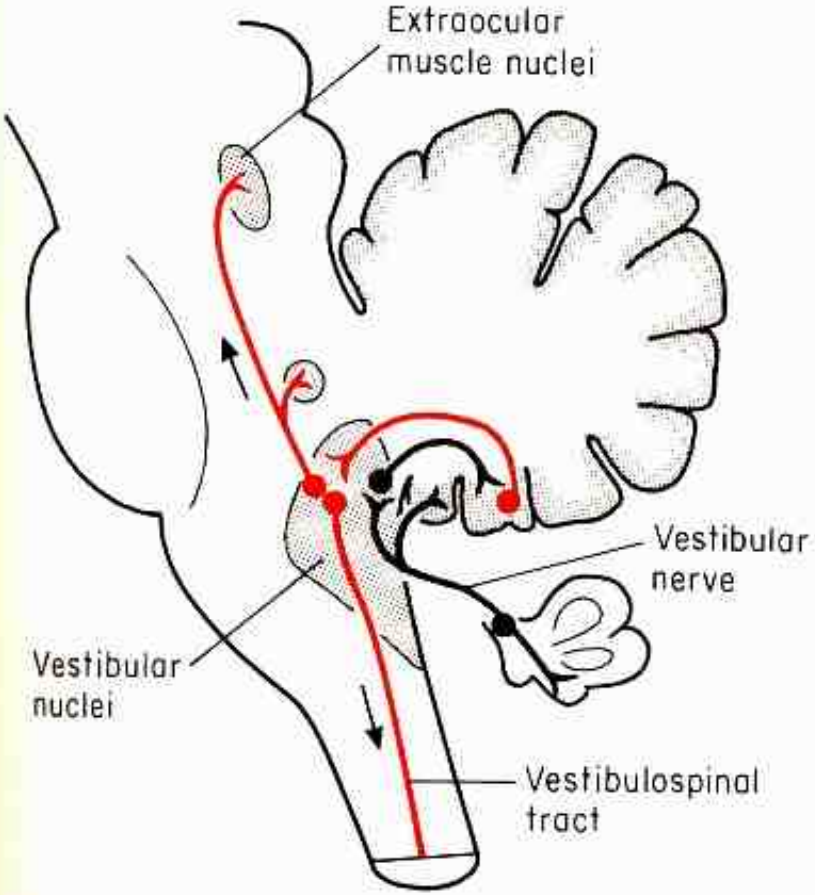


Fig. 11.3. *The main connections of the vestibulocerebellum.* Afferents are shown in black and efferents in red in a schematic drawing of a sagittal section through the brain stem. Note primary and secondary vestibulocerebellar fibers and the projection back to the vestibular nuclei. In addition to afferents from the vestibulocerebellum, the vestibular nuclei also receive cerebellar afferents from the anterior lobe vermis and from the fastigial nucleus.

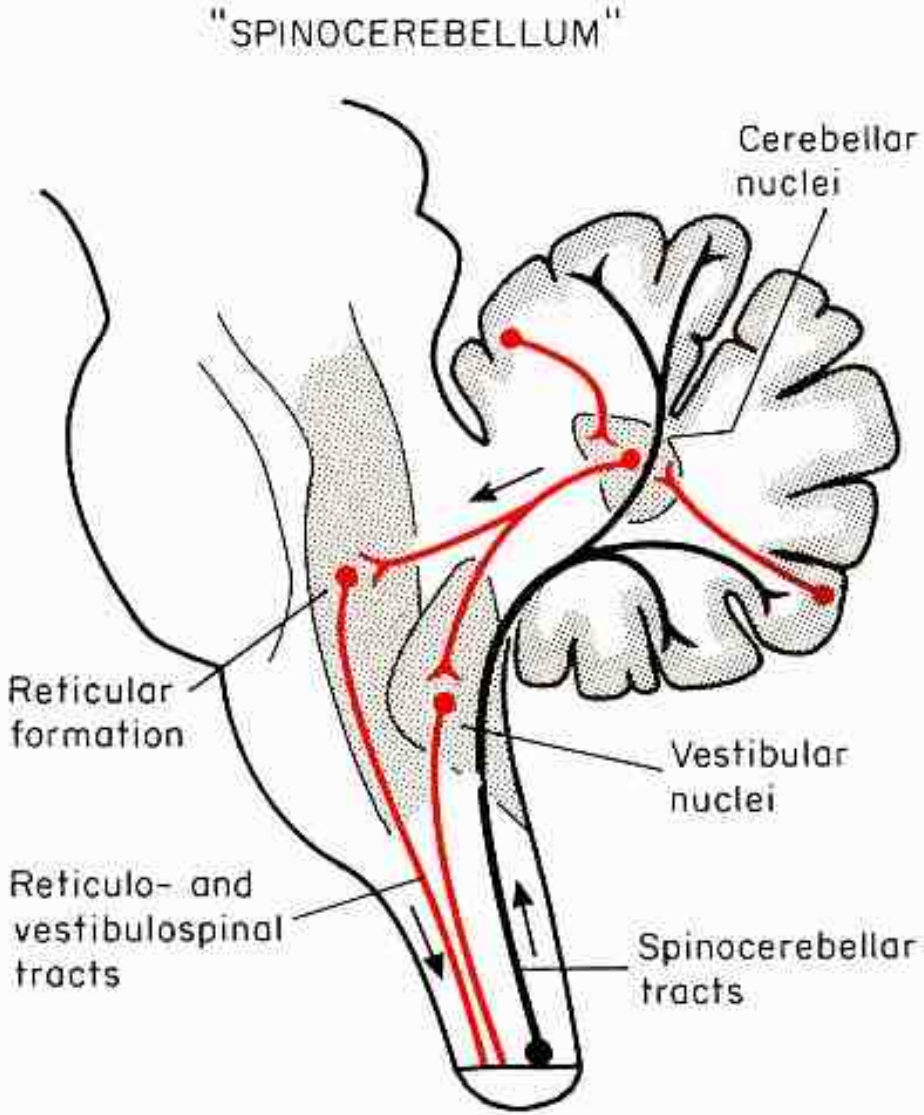


Fig. 11.4. *The main connections of the spinocerebellum.* Note that the spinocerebellum can influence spinal motoneurons via reticulospinal and vestibulospinal pathways.

CEREBELLUM

1) Receives extensive sensory input, but is not involved in voluntary discrimination or interpretation

2) Influences motor functions, but resection of the cerebellar cortex does not result in lasting paralysis

3) Stimulation of the cerebellar cortex does not evoke movements

4) Inflences cognitive functions namely motor learning and higher mental functions

Mozečkové syndromy

- **Flokulonodulární sy –** poruchy rovnováhy, nejistota při chůzi a stoji, nystagmus
- **Sy předního laloku –** nejistá chůze (ataxie, zejména při lézi vermis a paravermální oblasti), alkohol
- **Neocerebellární sy** – koordinace volních pohybů, poh\b\ nejisté, trhané, neplynulé, dekomposice pohybů, dysmetrie (pohyb není včas zastaven) dysdiadochokinesa, intenční třes, mozečková řeč,
- Kognitivní funkce

Paleocerebellar lesions (syndrome)

MIDLINE LESIONS

The midline portions of the cerebellum may be invaded by a tumor, typically a "medulloblastoma" that occurs in childhood. In adults, a similar syndrome may be seen in chronic alcoholism, which causes degeneration of the vermis. The patient has an unsteady, staggering **ataxic gait**, walks on a wide base, and sways from side to side. **Cerebellar nystagmus** is "pendular," with eye movements of equal speed in both directions, usually in the horizontal plane. It is attributed to interruption of connections of the vermis with the ocular motor nuclei by way of the vestibular nuclei and the reticular formation. The signs are at first limited to a disturbance of equilibrium; however, additional cerebellar signs appear when a tumor invades other parts of the cerebellum.

NEOCEREBELLAR SYNDROME (LESION OF THE HEMISPHERE)

The following signs, in varying degrees of severity, are those of a neocerebellar syndrome. Movements are **ataxic** (intermittent or jerky). There is **dysmetria**; for example, when the patient reaches out with the finger to an object, the finger overshoots the mark or deviates from it (**past-pointing**). Rapidly alternating movements, such as flexion and extension of the fingers or pronation and supination of the forearm, are performed in a clumsy manner (**adiadochokinesis**). **Asynergy** is separation of smoothly flowing voluntary movements into successions of mechanical or puppet-like movements (**decomposition of movement**). There may be **hypotonia** of muscles, which also tire easily. Cerebellar **tremor**, which occurs most frequently with demyelinating lesions in the cerebellar peduncles, usually occurs at the end of a particular movement (**intention tremor**). **Dysarthria** is evident if asynergy involves muscles used in speech, which is then thick and monotonous (slurring; scanning speech). There may be nystagmus, if the lesion encroaches on the vermis. The deficits noted are superimposed

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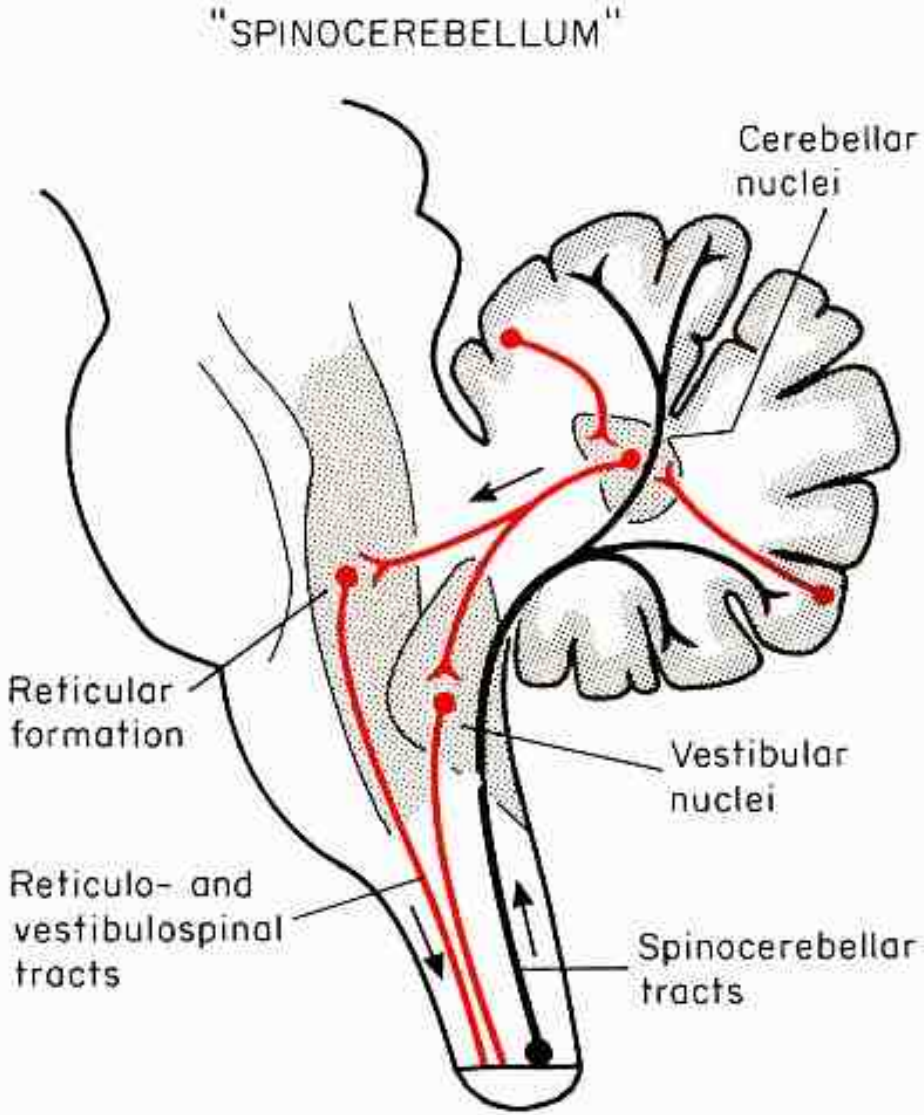


Fig. 11.4. *The main connections of the spinocerebellum.* Note that the spinocerebellum can influence spinal motoneurons via reticulospinal and vestibulospinal pathways.

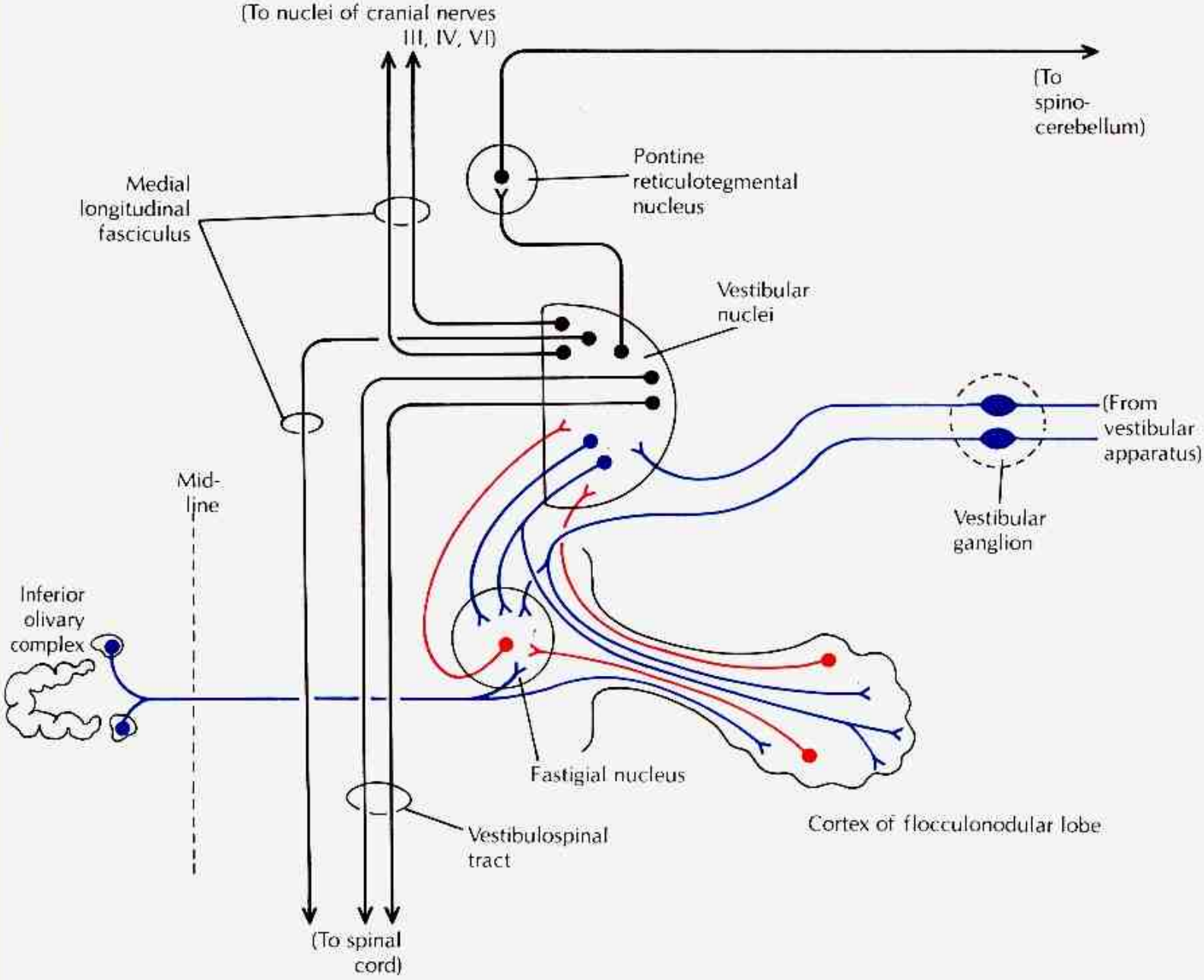


Figure 10-13. Connections of the vestibulocerebellum and vestibular nuclei.

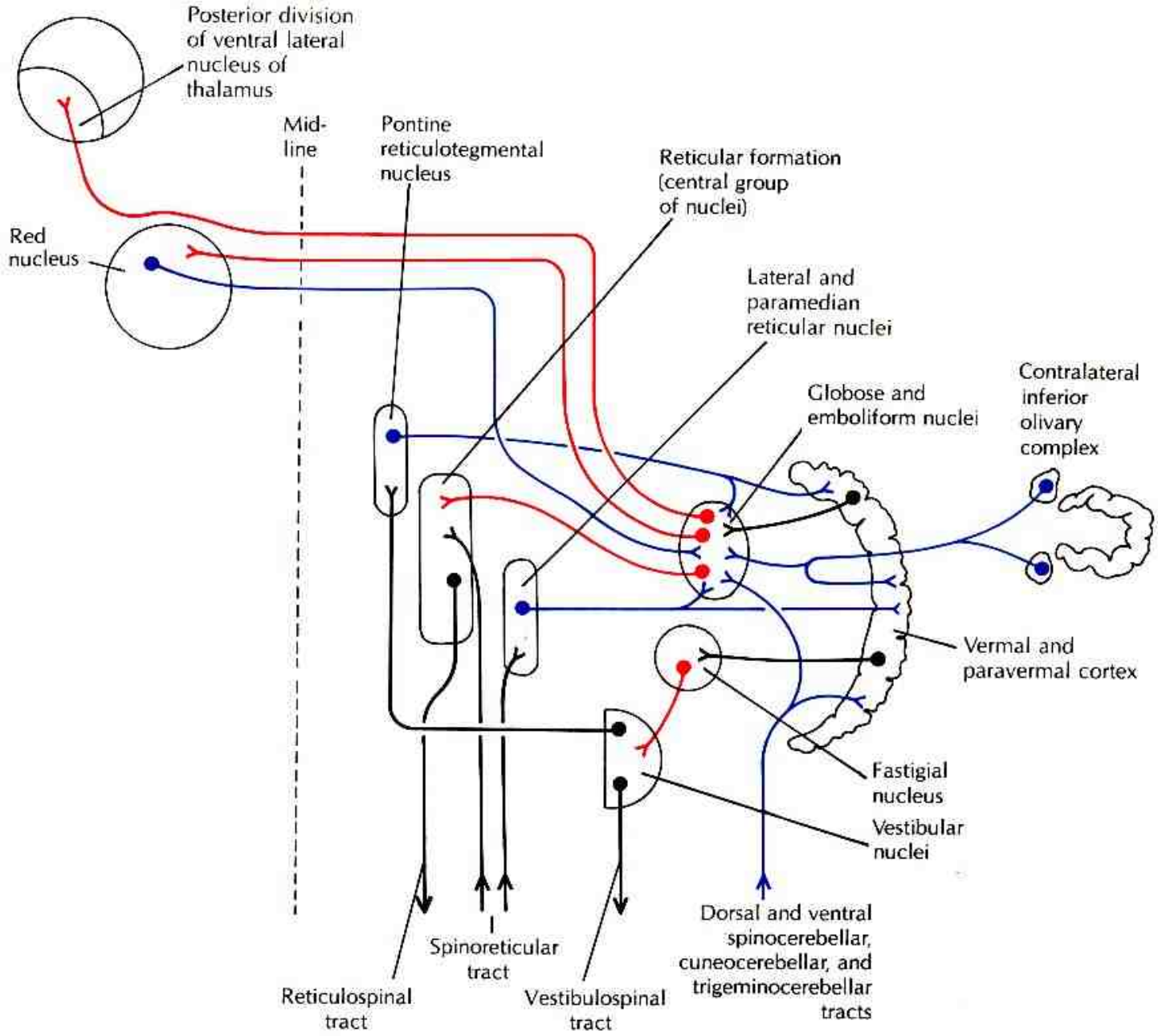


Figure 10-14. Connections of the spinocerebellum.

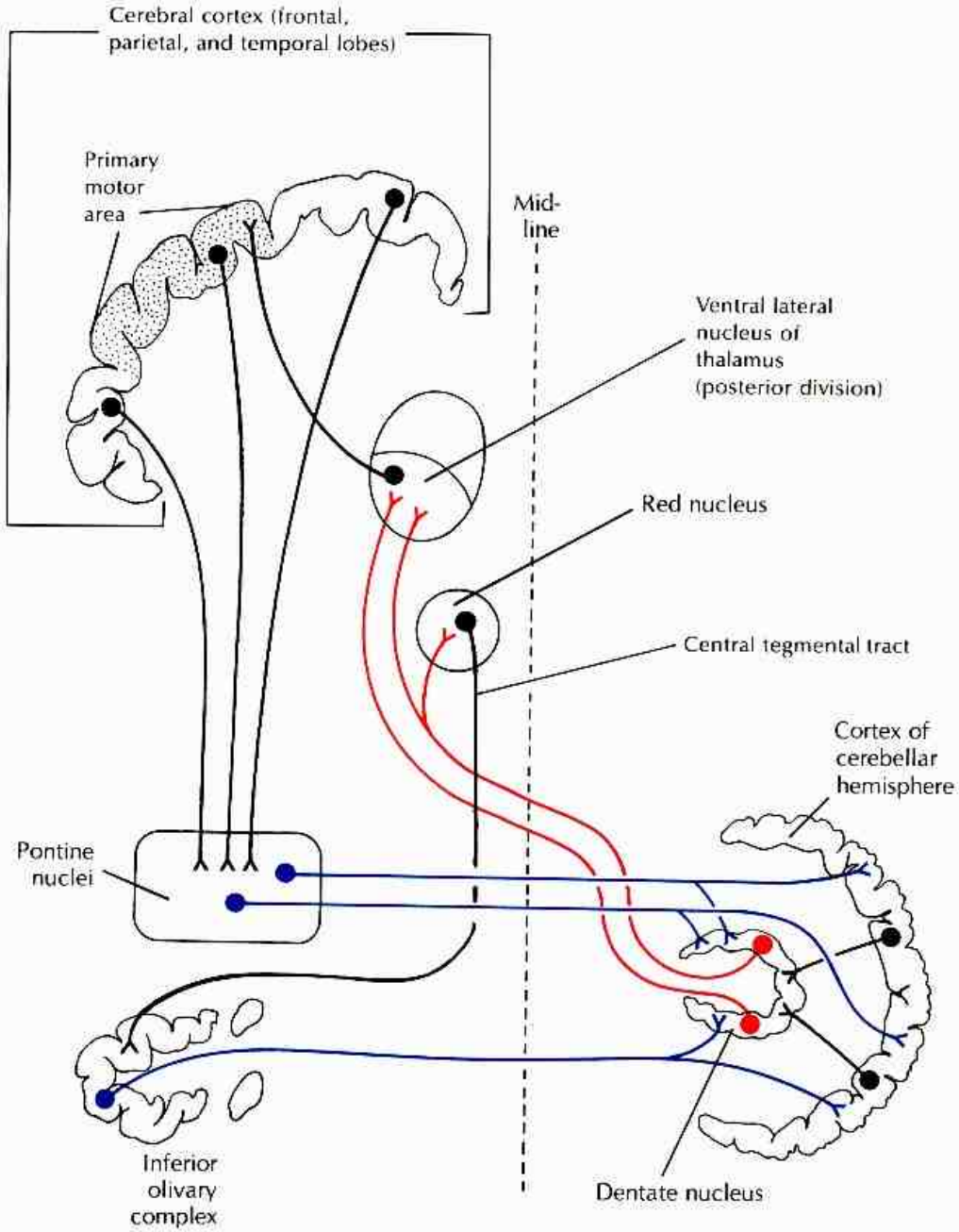


Figure 10-15. Connections of the pontocerebellum.

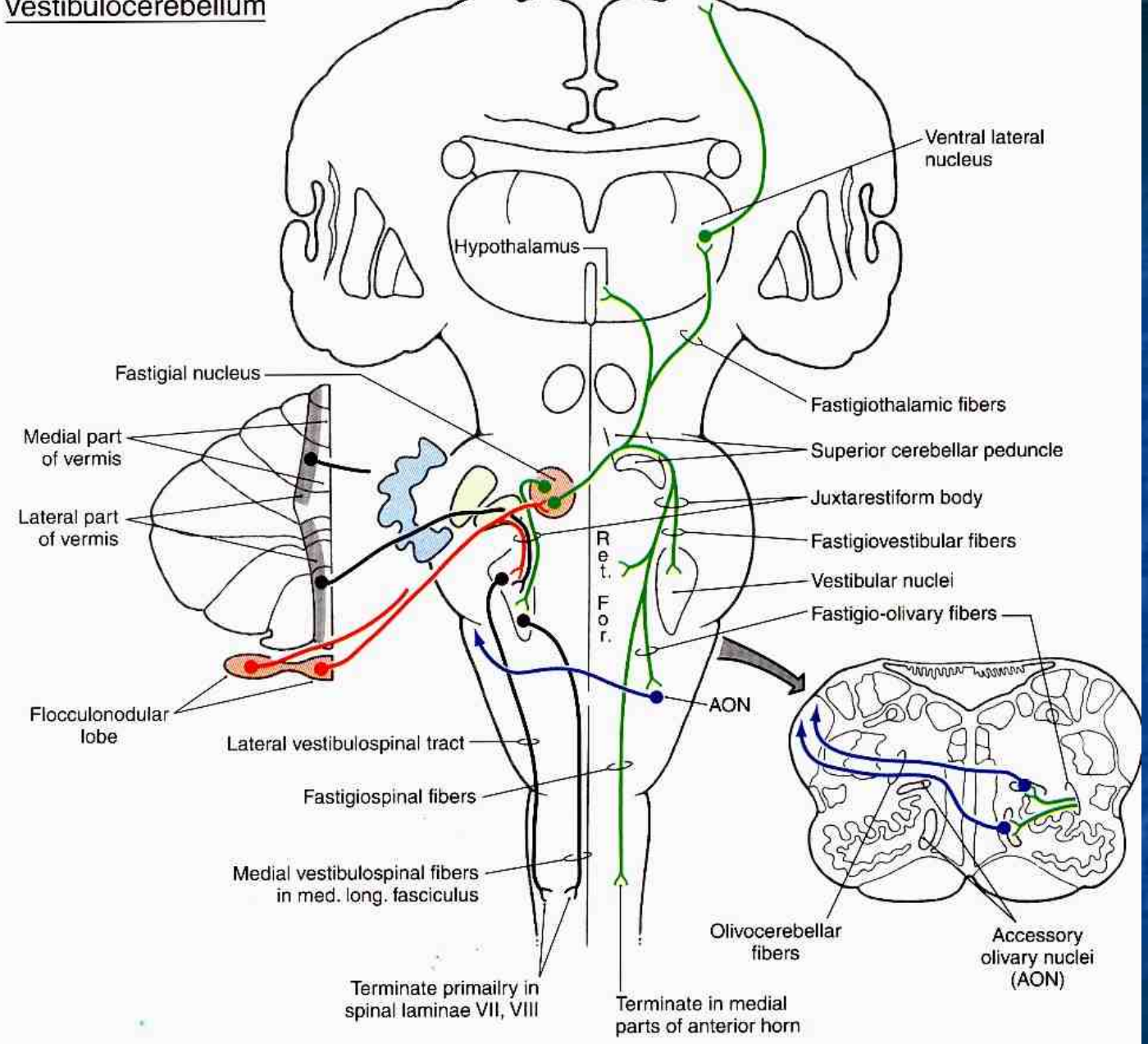


Figure 27–14. Projections of the vestibulocerebellum and of the lateral part of the medial zone through the fastigia vestibular nuclei. med. long., medial longitudinal; Ret. For., reticular formation.

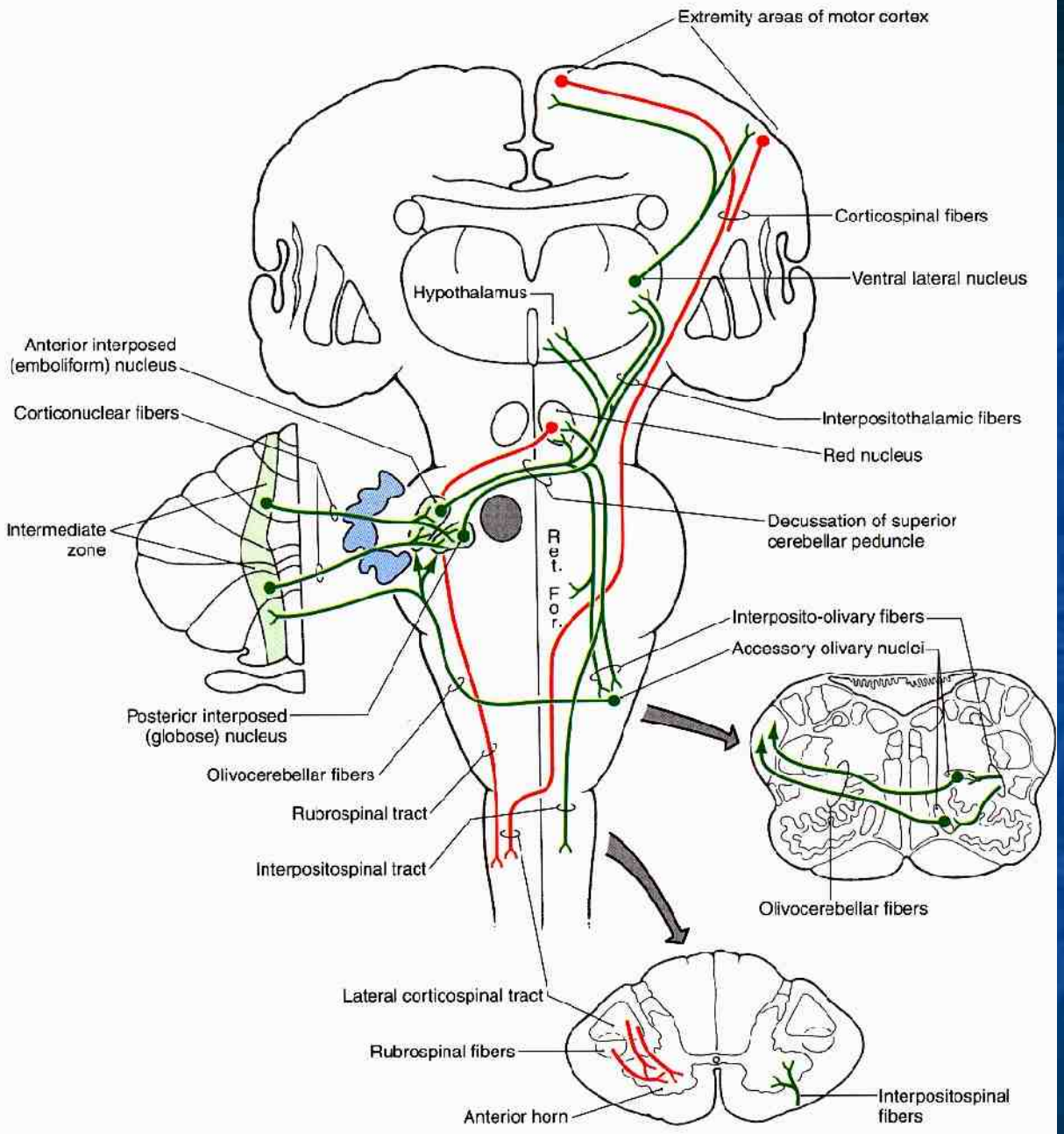


Figure 27–16. Projections of the spinocerebellum (intermediate zone) through the emboliform and globose nuclei. Ret. reticular formation.

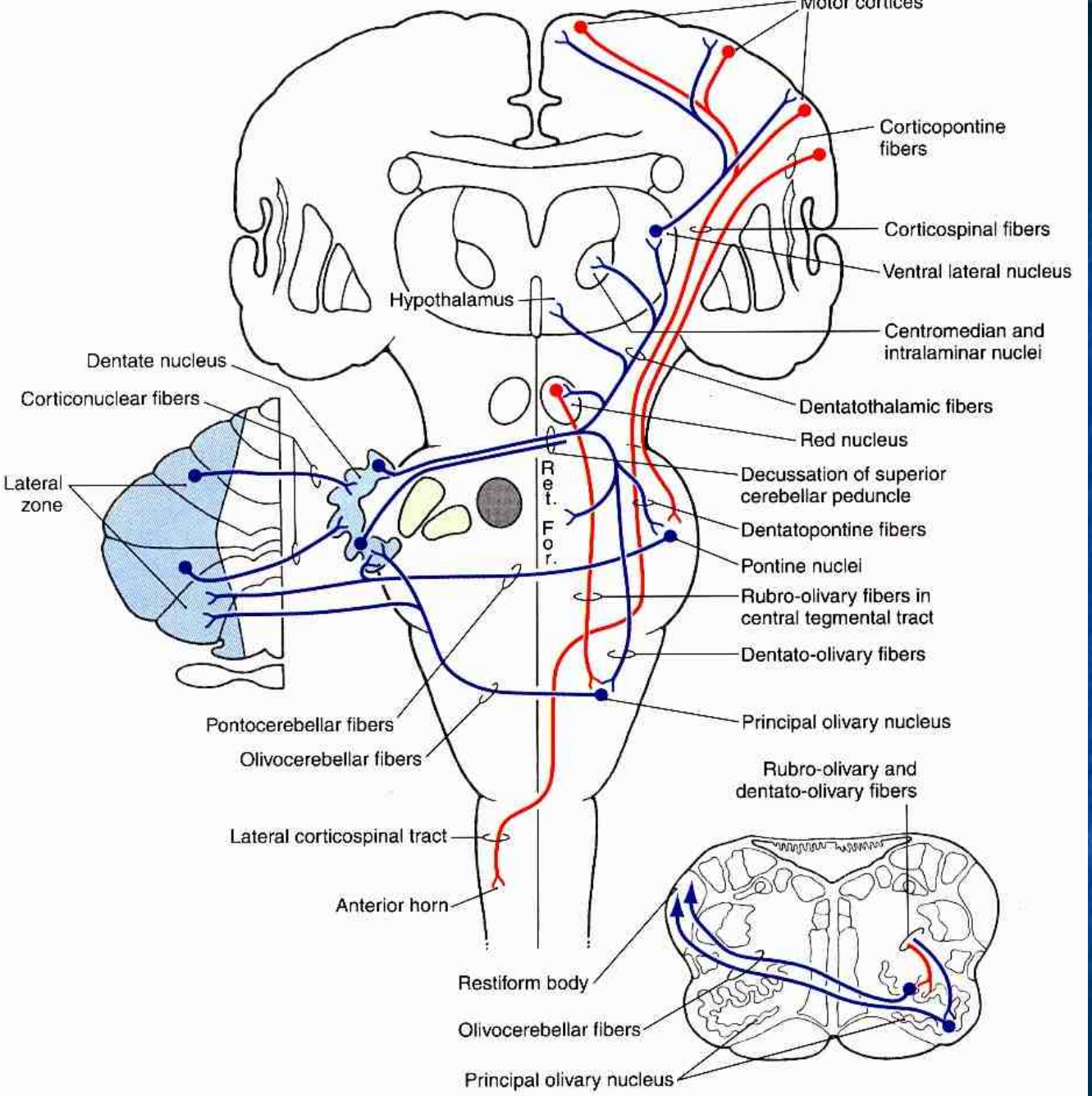
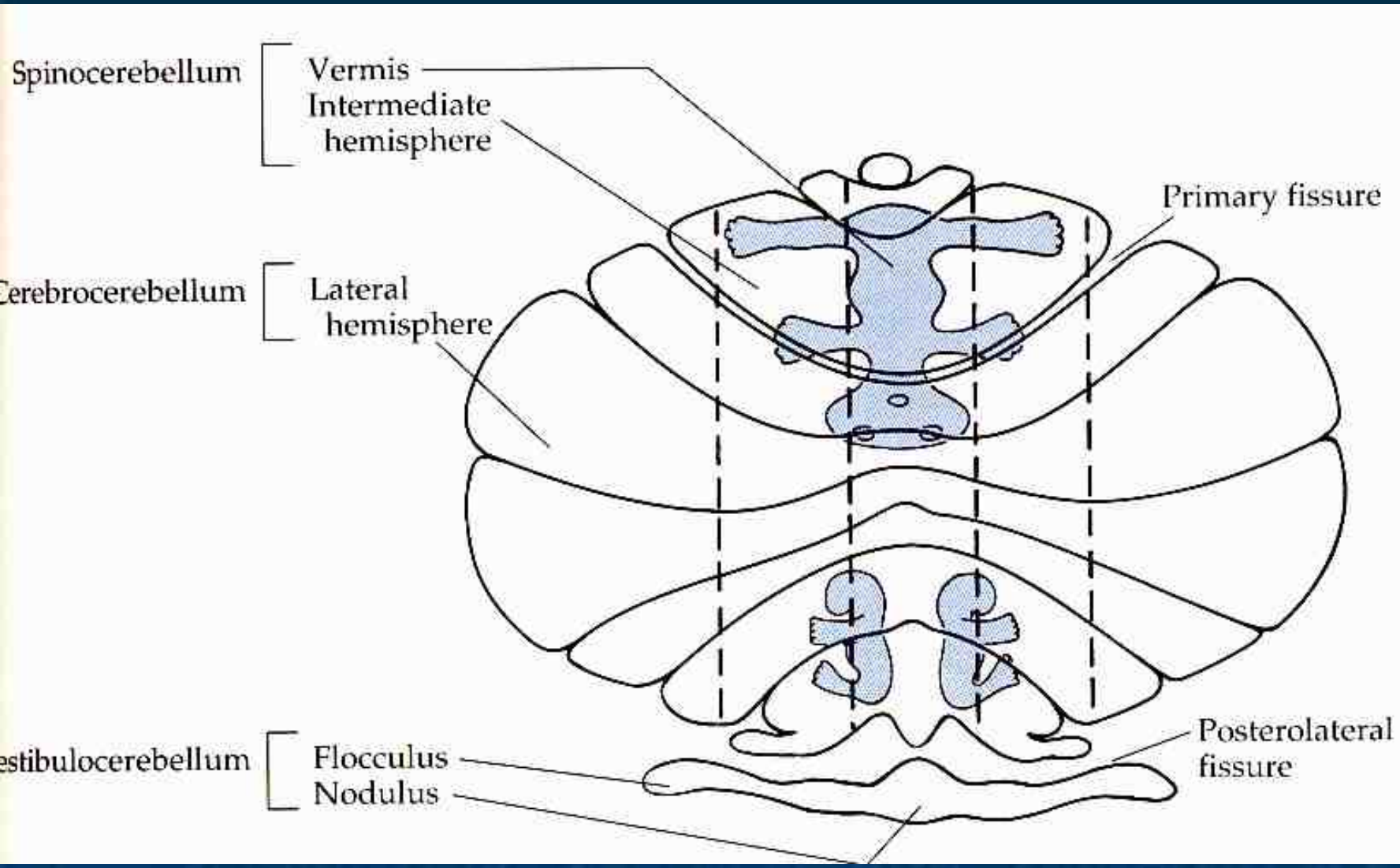
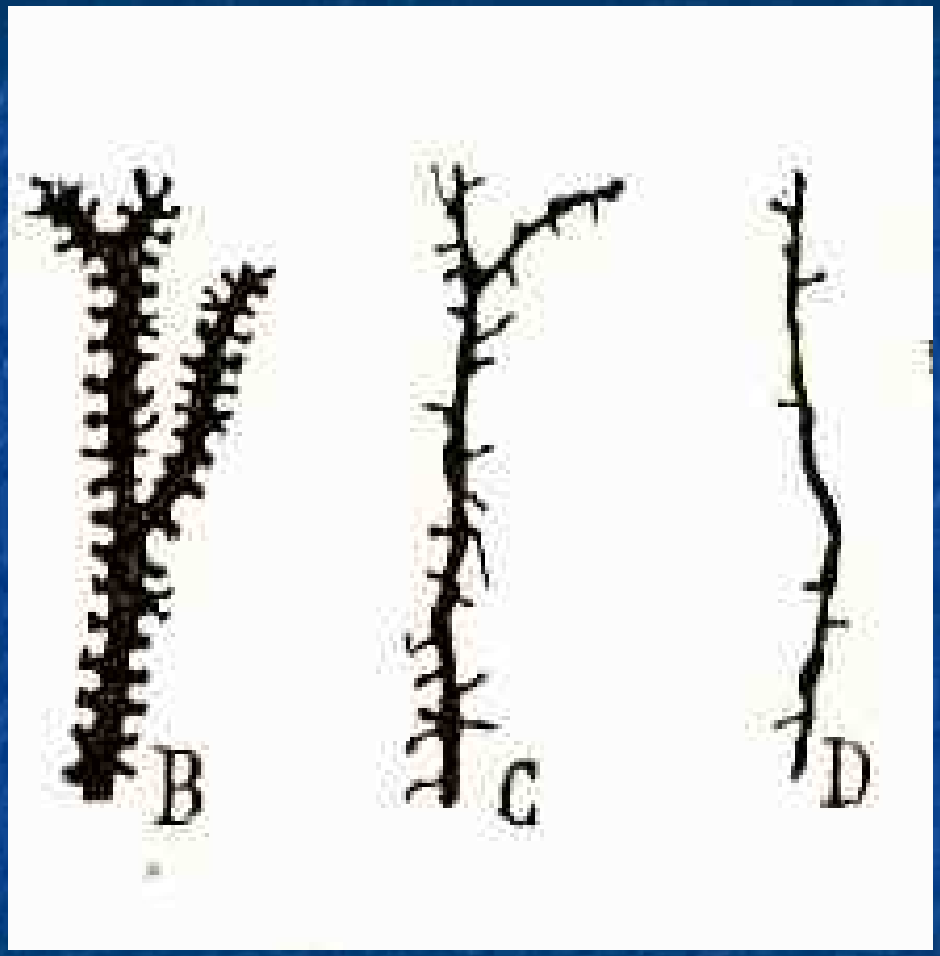


Figure 27–17. Projections of the pontocerebellum (lateral zone) through the dentate nucleus. Ret. For., reticular forma


Spinocerebellum
Vermis
Intermediate hemisphere
Primary fissure
Cerebrocerebellum
Lateral hemisphere
Vestibulocerebellum
Flocculus
Nodulus
Posterolateral fissure


B
C
D

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Cajal 1911

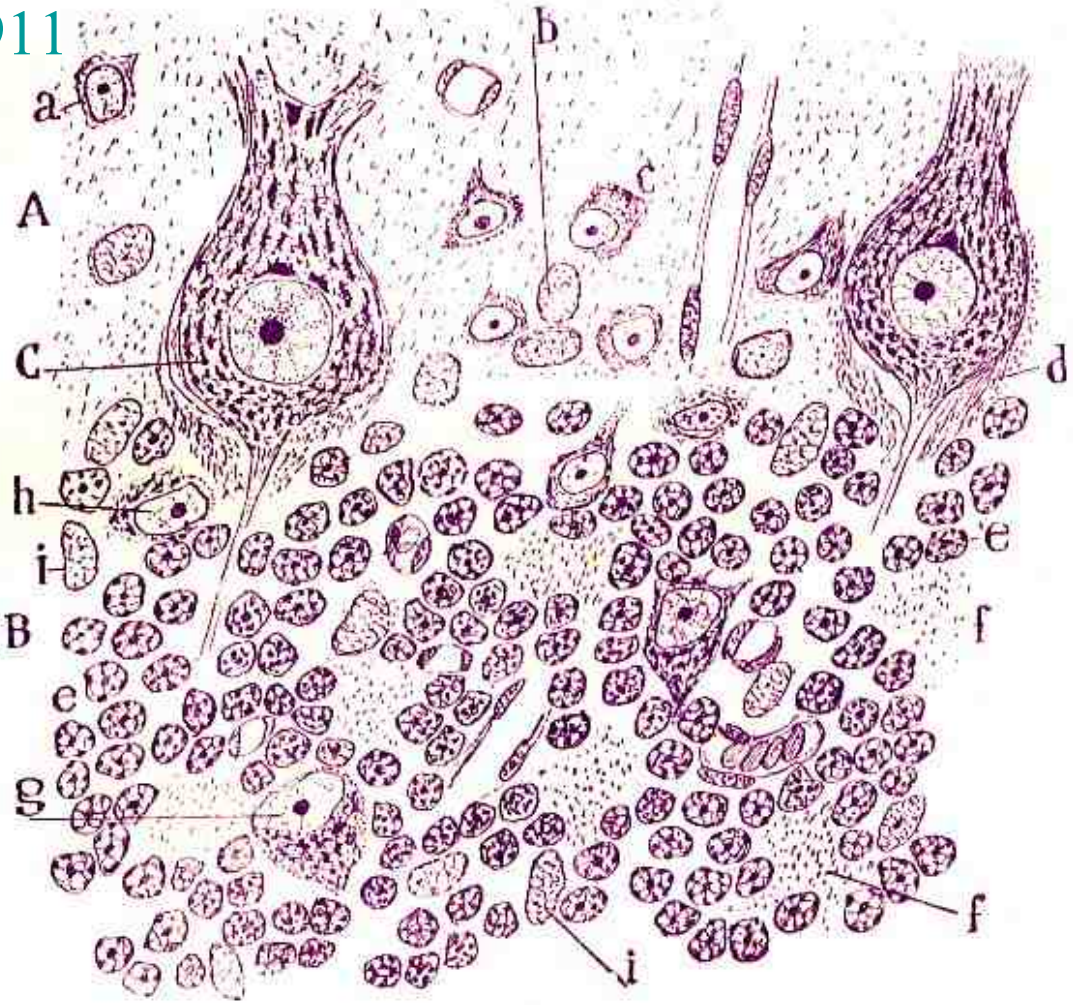

FIG. 1. — Portion d'une coupe de l'écorce du cervelet ; homme adulte. Méthode de Nissl ; obj. apochrom. 1,30.

A, région inférieure de la couche plexiforme ; — B, couche des grains ; — C, corps des cellules de Purkinje ; — *a*, cellule étoilée de la couche plexiforme ; — *b*, noyaux des cellules épithéliales ; — *c*, autre cellule étoilée avec chromatine marginale ; — *d*, masse fibrillaire correspondant aux corbeilles qui embrassent les corps des cellules de Purkinje ; — *e*, noyaux des grains ; — *f*, îlots granuleux ou cérébelleux ; — *g*, *h*, cellules de Golgi ou à cylindre-axe court de la zone des grains ; — *i*, noyaux des cellules névrogliques.

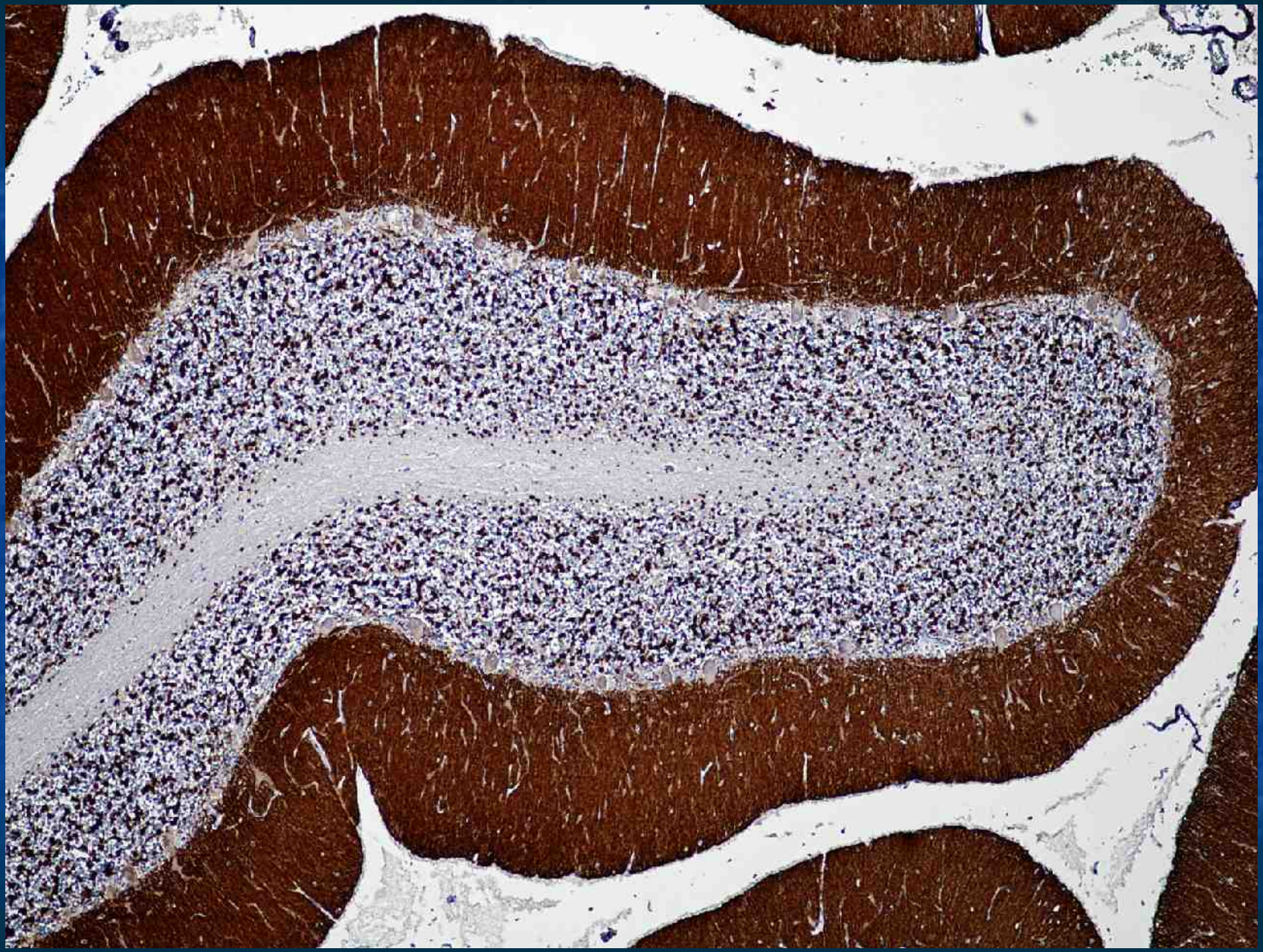

Image266.x4 synapto