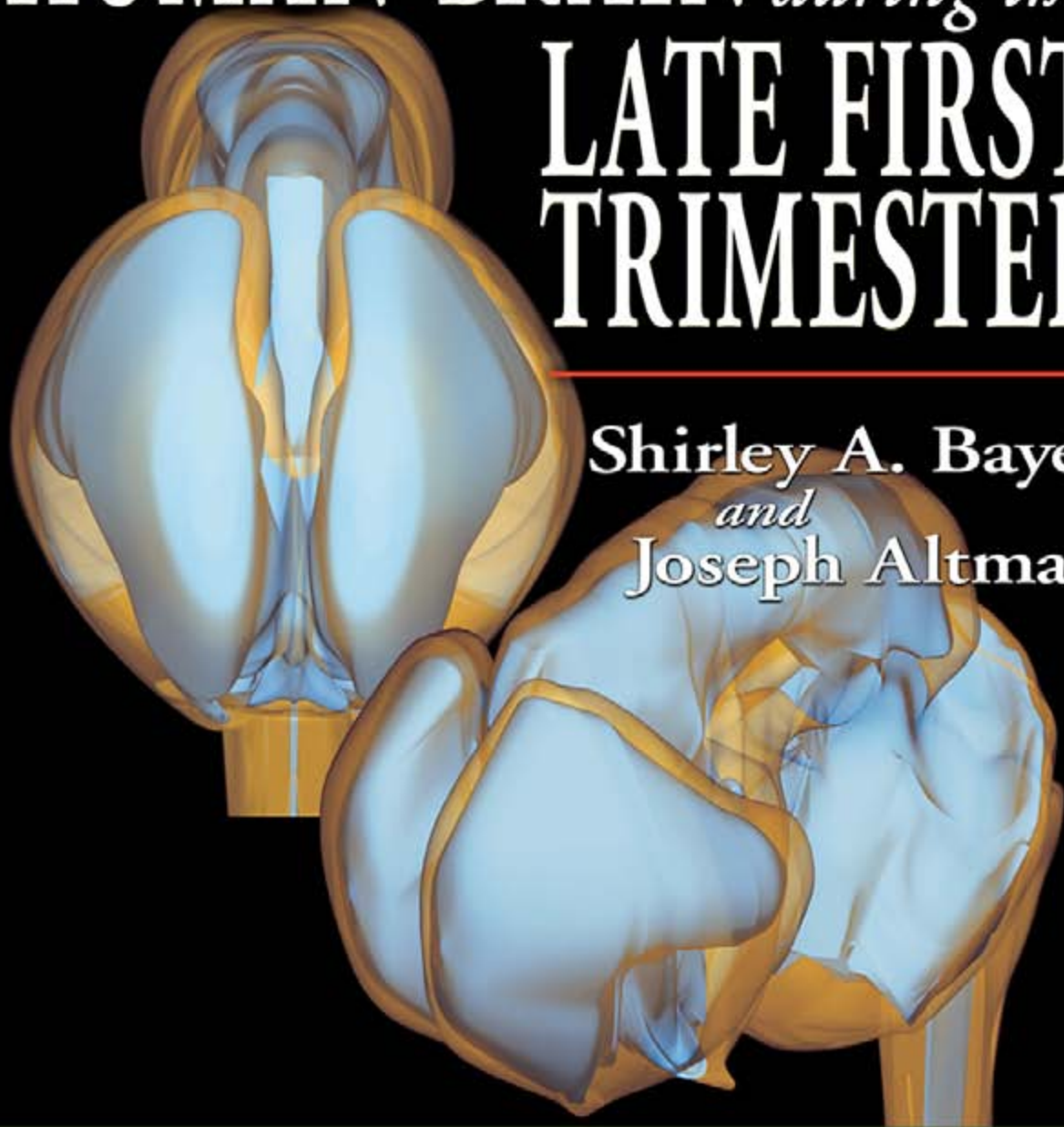


ATLAS OF
HUMAN CENTRAL NERVOUS SYSTEM DEVELOPMENT
VOLUME 4

The

HUMAN BRAIN *during the*
**LATE FIRST
TRIMESTER**

Shirley A. Bayer
and
Joseph Altman



Taylor & Francis
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HUMAN BRAIN *during the*
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**ATLAS OF
HUMAN CENTRAL NERVOUS SYSTEM DEVELOPMENT
SERIES**

Shirley A. Bayer *and* Joseph Altman

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The Spinal Cord from Gestational Week 4 to the 4th Postnatal Month

VOLUME 2

The Human Brain During the Third Trimester

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The
HUMAN BRAIN *during the*
LATE FIRST
TRIMESTER

Shirley A. Bayer and Joseph Altman



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Boca Raton London New York

CRC is an imprint of the Taylor & Francis Group,
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Published in 2006 by
CRC Press
Taylor & Francis Group
6000 Broken Sound Parkway NW, Suite 300
Boca Raton, FL 33487-2742

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CRC Press is an imprint of Taylor & Francis Group

No claim to original U.S. Government works
Printed in the United States of America on acid-free paper
10 9 8 7 6 5 4 3 2 1

International Standard Book Number-10: 0-8493-1423-2 (Hardcover)
International Standard Book Number-13: 978-0-8493-1423-0 (Hardcover)

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Library of Congress Cataloging-in-Publication Data

Catalog record is available from the Library of Congress

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DEDICATION

We dedicate this volume to the memory of Prof. Dr. Ferdinand Hochstetter (1861-1954) for his pioneering description of the development of the human brain during the first trimester. To this day, his publication that appeared in Vienna and Leipzig in 1919 is the best source for systematically showing the external features of the human brain at this stage, and the current work is built on the foundation he provided.

ACKNOWLEDGMENTS

We thank Dr. William DeMyer, pediatric neurologist at Indiana University Medical Center, for access to his personal library on human central nervous system development. We also thank the staff of the National Museum of Health and Medicine at the Armed Forces Institute of Pathology, Walter Reed Hospital, Washington, D.C.: Dr. Adrienne Noe, Director; Archibald J. Fobbs, Curator of the Yakovlev Collection; Elizabeth C. Lockett; and William Discher. We are most grateful to Dr. James M. Petras at the Walter Reed Institute of Research who made his darkroom facilities available so that we could develop all the photomicrographs on location rather than in our laboratory in Indiana. Finally, we thank publisher Barbara Norwitz, project manager Jim McGovern, proofreader Samar Haddad, Randy Brehm, and Kari Budyk at CRC Press/Taylor and Francis for their personal attention to us and for expert help during production of the manuscript.

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PART I

INTRODUCTION

A. Organization of the Atlas

This Atlas focuses on the development of the human brain during the late first trimester, and is Volume 4 in the *Atlas of Human Central Nervous System Development* series. Volume 1 (Bayer and Altman, 2002) provides a record of the development of the spinal cord from the 4th gestational week (GW4) to the 4th postnatal month. Volume 2 (Bayer and Altman, 2004a) records brain development during the third trimester, with the specimens ranging in age from GW37 to GW26. Volume 3 (Bayer and Altman, 2005) presents brain development during the second trimester, from GW24 to GW13.5. The specimens dealt with in the present volume cover the period from GW11 to GW7.5. The major theme of Volume 2 is the *maturation* of the brain's *settled* and *enduring* neuron populations. The major theme of Volume 3 is the *migration, sojourning, and settling* of neuronal populations. Prominent migratory streams and sojourning structures are evident by GW13.5 and many of these are still present at GW24. The major theme of the present volume is *late neurogenesis* and *early neuronal differentiation*. The structures of central interest are the visually distinguishable divisions, or mosaics, of the germinal neuroepithelium that generate different populations of neurons and neuroglia. The major theme of the forthcoming Volume 5, which will deal with brain development during the early first trimester (GW3-GW7), will be *neuroepithelial expansion* along the *expanding brain ventricles*, and *early neurogenesis*. As in Volumes 2 and 3, the specimens are presented in a reverse, older-to-younger order. We do that for both heuristic and pedagogical reasons: proceeding from the familiar and better known to what is less familiar and often uncertain or hypothetical.

The present volume contains grayscale photographs of nine sets of Nissl-stained, and one set of Bodian-stained sections, of normal brains ranging in age from early fetuses,

crown-rump (CR) length 57-60 mm and estimated age GW11, to late embryos, CR length 21-23 mm and estimated age GW7.5. As in the previous volumes, we sought to select for each age group, brains sectioned in the coronal, horizontal, and sagittal planes. However, this posed a problem as the terms “coronal” and “horizontal” (but not “sagittal”) become more and more ambiguous when applied to younger and younger fetal brains and then embryonic brains. The conventional neuroanatomical designation of “coronal” comes from the practice of placing the dissected brain on a horizontal surface, and with the base of the cerebral cortex and the cerebellar cortex lying flat on that surface, making transverse (left-to-right) cuts perpendicular to the supporting surface. In a similar manner, “horizontal” sections are made by making the cuts parallel to the surface. With this convention, there is little ambiguity, for instance, that the frontal lobe is situated anteriorly and the occipital lobe posteriorly. However, this practice has not been consistently followed by embryologists; indeed, it could not be followed when the brains are not shelled out and the embryo is sectioned *de toto*. Moreover, since the whole embryo, and more particularly the medulla, pons and brainstem (as described in this volume) undergo a series of flexures, a coronal cut made at one point becomes an oblique or horizontal cut at another point. The same applies to horizontal cuts. Indeed, perusal of embryological textbooks reveals little consistency in the positioning of an embryonic brain or series of brains in the illustrations. In order to produce consistency in presenting the human brains of different fetal and embryonic ages, and facilitate comparisons of the changes across all the ages, we have therefore adopted the following procedure. Each brain, irrespective whether it was designated in the original protocol as “coronal” or “horizontal,” has been placed into an X-Y coordinate frame as if it were a mature brain; i.e., the ventral surface of the cortex and the cerebellum lie flat on an imaginary line that we call the “cardinal horizontal

plane.” The line perpendicular to it, we call the “cardinal coronal plane.” Given this coordinate system we then indicate the deviation of a given brain from the cardinal coronal or cardinal horizontal. This method allows a consistent positioning of the telencephalon, diencephalon, mesencephalon, and cerebellum (but not of the pons, medulla and spinal cord) into an age-independent stereotactic framework, and designations like anterior and posterior, and dorsal and ventral pointing in the same direction for easy comparisons across ages.

Each specimen is presented in a separate part of the Atlas: GW11 in the coronal plane in **Part II**; GW11 in the horizontal plane in **Part III**; GW9 in the sagittal plane in **Part IV**; GW9 in the horizontal plane in **Part V**; GW9 in the coronal plane in **Part VI**; GW8 in the sagittal plane in **Part VII**; GW8 in the coronal plane in **Part VIII**; GW8 in the horizontal plane in **Part IX**; GW7.5 in the coronal plane in **Part X**; and GW7.5 in the sagittal plane in **Part XI**. Selected “coronal” plates are presented from rostral to caudal in portrait orientation. The dorsal part of each section is toward the top of the page, the ventral part at the bottom, and the midline is in the vertical center of each section. Sagittal plates are presented from medial to lateral in portrait orientation. The anterior part of each section is facing left, posterior right, dorsal top, and ventral bottom. “Horizontal” plates are presented from rostral to caudal in landscape orientation. The anterior part of each section is facing to the left (bottom of page), posterior to the right (top of page), and the midline is in the horizontal center of each section. Each part contains *companion plates*, designated as **A** and **B** on facing pages; some of the plates are expanded into **C** and **D** parts. Parts **A** and **C** of each plate on the left page show the full contrast photograph without labels; parts **B** and **D** show low contrast copies of the same photograph on the right page with superimposed outlines of structures and unabbreviated labels. The *low magnification plates* show entire sections to identify the large structures of the brain, such as the various lobes and gyri of the cerebral cortex, and large subdivisions of the brain core, such as the basal ganglia, thalamus, hypothalamus, midbrain, pons, cerebellum, and medulla. The *high magnification plates* feature enlarged views of the brain core to identify smaller structures. For ease of interpretation in all plates, the ventricles are labeled in **CAPITALS**, the neuroepithelium and other germinal zones in **Helvetica bold**, transient structures in **Times bold italic**, and permanent structures in Times Roman or **Times bold**. Fixation artifacts and processing damage are usually outlined with *dashed lines* in parts **B** or **D** of each plate. Finally, an alphabetized **Glossary** gives brief definitions of most labels used in the plates with expanded definitions of all transient developmental structures.

B. Specimens

A total of 10 specimens are illustrated and annotated in this Volume. All of them are from the Collections of human embryos or fetal brains currently kept at the National Museum of Health and Medicine, Armed Forces Institute of Pathology, Washington, D.C. One brain, estimated age GW11, is from the Yakovlev Collection and is designated, using the prefix Y, as Y1-59. We give a brief description of the Yakovlev Collection in Volume 2 (Bayer and Altman, 2004a) and more detail is provided in Haleem (1990). Eight specimens are from the Carnegie Collection and are designated by their respective numbers with the added prefix C. The Carnegie Collection was started by Franklin P. Mall (1862-1917) and expanded at the Carnegie Institution of Washington under the direction of George L. Streeter (1873-1948) and George W. Corner (1889-1981). One specimen, estimated age GW9, is from the Minot Collection and is designated, using the prefix M, as M841. The Minot Collection is named after Charles S. Minot (1852-1914), who collected and prepared more than 1900 embryos of different animal species, and about 100 human embryos, close to a century ago. We made use of models of the embryonic and fetal brains prepared in the Carnegie Institution, and by Hochstetter (1919).

C. Photography and Computer Processing

Selected sections of the Yakovlev specimen (Y1-59) were photographed at low magnification, using a macro lens (Vivitar, 55mm 1:2.8 auto macro) attached to a Nikonmat 35mm camera that was mounted on a stand over a fluorescent light board. The Carnegie and Minot specimens and all higher magnification views were photographed using either an Olympus photomicroscope or a Wild phototmakroskop. The magnification varied for each specimen according to the size of the dissected brain or entire fetal head: the section with the largest area that could be accommodated within the field of view set the magnification for all sections of a particular specimen. All photographs were taken with a green filter to increase the contrast of the black and white film (Kodak technical pan #TP442). The film was developed at 20°C for 6 to 7 minutes in Kodak HC110 developer (dilution F), followed by Kodak stop bath for 30 seconds, Kodak fixer for 5 minutes, Kodak hypo clearing agent for 1 minute, running water rinse for 10 minutes, and a brief rinse in Kodak photoflo before drying.

The negatives were scanned at 2700 dots-per-inch (dpi) with a Nikon Coolscan-1000 35mm film scanner which was interfaced to a PowerPC G3 Macintosh computer running Adobe Photoshop (version 5.02) with a plug-in Nikon driver. To capture the subtle shades of gray, the negatives were scanned as color positives, inverted, and converted to grayscale. Using the enhancement features built into Adobe Photoshop and the additional features of Extensis

Intelligence, adjustments were made to increase contrast and sharpness. When the image resolution was set to 300 dpi, a full-size photographic file printed at approximately 12-10 inches. Images are shown at slightly reduced full size on separate pages. Adobe Illustrator was used to superimpose labels and outline structural details on low contrast copies of the Adobe Photoshop files. The Plates were placed into a book-form layout using Adobe InDesign. Finally, camera-ready files were provided to Taylor and Francis in Adobe portable document format.

The entire brain and upper cervical spinal cord of C966 was three-dimensionally reconstructed to show the large superventricles and the surface features of the neuroepithelium in the telencephalon, diencephalon, and rhombencephalon (**Figures 10-19, Part X**). The brain reconstructions (**Figures 10-15**) involved five steps. *First*, photographs of serial sections were made throughout the entire brain; the negatives were scanned and converted to computer files as described in the preceding paragraph. *Second*, all the files of sections selected for the reconstruction were placed into one large Photoshop file that contained a separate photograph in each layer. By altering the visibility and transparency of these layers the sections were aligned to each other as they were before sectioning. Then each layer was saved as a separate file. *Third*, Adobe Illustrator was used to outline the brain surface and the edge of the ventricles, and the outlines of each section were saved in separate Adobe Illustrator eps (encapsulated postscript) files. *Fourth*, the eps files were imported into 3D space (x, y, and z coordinates) using Cinema 4DXL (C4D, Maxon Computer, Inc.), a modeling and animation software package. For each section, points on the outlines have unique x-y coordinates and the same z coordinate. By calculating the distance between sections, the entire array of outlines was stretched out in the z axis. The outlines were segregated into two groups, one for the brain surface, the other for the ventricular surface. The C4D loft tool builds a "skin" for each group of outlines by creating a spline mesh of polygons. The polygons start from the x-y points on the first outline with the most anterior z coordinate, to the x-y points on the next outline behind it, and finish with the x-y points on the last outline at the most posterior z coordinate. The spline meshes were rendered either as completely opaque or partly transparent surfaces using the C4D ray-tracing engine. Selected surfaces can be made either invisible or visible using the various options in C4D. The complex structure of the brain requires that the surface be built in several different lofts to avoid twisting the surface in the regions of the brain flexures. When these different loft segments are shown together, there are a few unavoidable artifacts, such as surface indentations and changes in the way light reflects from the surface; some of these are labeled in **Figures 10-19**. *Fifth*, the rendered images were converted to Adobe Photoshop files, and Adobe Illustrator was used to label the structures and to draw thin lines on some of the surfaces to make the images easier to under-

stand. The reconstruction of the parts of the neuroepithelium (**Figures 16-19**) followed the same five steps except that only the selected parts of the neuroepithelium in each section were outlined, and these were always rendered as completely opaque blocks of tissue.

D. Identification of Immature Brain Structures

In contrast to the mature human brain, there are no comprehensive textbooks or atlases available on the entire course of the human brain development. The few publications that cover the late first trimester give overviews that identify the largest brain structures in chapters included in embryology textbooks (e.g., Patten, 1953; Hamilton et al., 1964) and atlases (Gasser, 1975; O'Rahilly and Müller, 1994). Some facets of human brain development during this period are part of an overview in an edited publication (Sidman and Rakic, 1982), and there are a few research papers available on the development of specific brain regions (e.g., Gilbert, 1935; Pearson, 1941; Kappers, 1958; Humphrey, 1960, 1968; Richter, 1965; Kahle, 1956; Hewitt, 1958; Shuangshoti and Netsky, 1966; Rakic and Sidman, 1970; Moore et al., 1999; Forutan et al., 2001; Koutcherov et al., 2002). There is no single publication that labels both large and small structures in the developing human brain in serially sectioned specimens at several different ages during the late first trimester. Many of the transient features of the early fetal/late embryonic human brain have never been studied. Therefore, we rely to a great extent on our experimental work in the developing rat brain to identify formative brain structures, migratory streams, and components of the germinal neuroepithelium. In those studies, we labeled proliferating cells in the germinal matrices of the brain with ^3H -thymidine administered at daily intervals through the entire prenatal period, and every other day up to weaning. By varying survival times after ^3H -thymidine exposure from 1 hour, to days, we used the autoradiographic technique to establish quantitative time-tables of neurogenesis, trace the speed and route of neuronal migrations, and document the settling patterns of neurons in the maturing brain. The results of these studies were published over a period of three decades in journal articles (see below) and reviewed in chapters contributed to edited books (Altman and Bayer, 1975, 1995, 2004; Altman, 1992; Bayer and Altman, 1995a, 1995b). We have also presented evidence for many parallels in the sequential order of brain development in rat and man (Bayer et al., 1993). Readers interested in details of those experimental results and the rationale of some of the anatomic and morphogenetic identifications made in this Atlas may consult the following publications.

Amygdala: Bayer (1980c).

Basal Ganglia: Bayer (1984, 1985b, 1987).

Cerebellum: Altman and Bayer (1978a, 1982, 1985a, 1985b, 1985c, 1997).

Cerebral Cortex: Altman and Bayer (1990a, 1990b); Bayer and Altman (1990, 1991a, Bayer et al., 1991a).
Cranial Nerve Nuclei: Altman and Bayer (1980a, 1980b, 1980c, 1982b).
Hippocampus: Altman and Bayer (1975, 1990c, 1990d, 1990e); Bayer (1980a, 1980b).
Hypothalamus: Altman and Bayer (1978c, 1978d, 1978e, 1986).
Medulla: Altman and Bayer (1980a, 1980b, 1980c, 1982b).
Midbrain: Altman and Bayer (1981a, 1981b, 1981c).
Olfactory Bulb: Altman (1969); Bayer (1983).
Pontine Area: Altman and Bayer (1978b, 1980d, 1987a, 1987b, 1987c, 1987d).
Preoptic Area: Bayer and Altman (1987).
Rhinencephalon: Bayer (1985a, 1986a, 1986b); Bayer and Altman (1991b).
Septal Area: Bayer (1979a, 1979b).
Spinal Cord: Altman and Bayer (1984, 2001).
Thalamus: Altman and Bayer (1979a, 1979b, 1979c, 1988a, 1988b, 1988c, 1989a, 1989b, 1989c).

E. Major Transitional Brain Structures in the Late First Trimester

Transitional brain structures in late first trimester fetuses include the following: (i) The neuroepithelium (NEP), the primary germinal matrix of neural stem cells that are the ultimate source of all the neurons and neuroglia of the central nervous system. (ii) The rhombencephalic, mesencephalic, diencephalic, and telencephalic superventricles that provide the greatly expanded cerebrospinal fluid (CSF) interface for the shuttling nuclei of the pseudostratified NEP cells that undergo mitotic division near the lumen. (iii) The expanding fetal choroid plexus (CP), and its later shrinkage and transformations, which appears to play a role in neurogenesis and neuronal differentiation. (iv) The spreading gliopithelia (GEP) along several fiber tracts, and the gliopithelia associated with the formative ependymal layer that lines the enduring portions of the brain ventricles (G/EP). (v) The secondary germinal matrices abutting the neuroepithelium, such as the subventricular zone (SVZ) of the cerebral cortex, and others situated some distance from the ventricles. (vi) Short-distance migratory routes, and long-distance migratory streams, such as the telencephalic rostral migratory and lateral migratory streams and several precerebellar migratory streams. (vii) The transient sojourn zones and stratified transitional fields, such as the cortical stratified transitional field (STF) and the cerebellar transitional field (CTF).

The Neuroepithelium and its Mosaic Organization. The NEP is a cell-dense pseudostratified proliferative matrix that lines the extensive ventricular system of the developing brain and spinal cord. It used to be called the ependymal layer, but that term is no longer used because the ependyma is a highly specialized tissue that lines the

shrunken ventricles of the maturing and adult central nervous system. The NEP is composed of stem cells that either differentiate as specialized populations of neurons and neuroglia or give rise to secondary germinal matrices some distance from the ventricular system. The latter include the SVZ of the cerebral cortex, the external germinal layer of the cerebellar cortex, and the subgranular zone of the hippocampal dentate gyrus. At the outset of its development, the NEP stem cells are the sole constituents of the brain (to be described and illustrated in Volume 5). The bulk of NEP cells either keep proliferating or generate differentiating (postmitotic) neurons that, following a precise spatio-temporal order, migrate over a short or long distance to form various brain structures. While NEP cells all look alike in Nissl-stained preparations, discontinuous NEP “stretches” and “patches,” or *mosaics*, can be distinguished along the ventricles in terms of matrix thickness (pseudostratified cell depth), cell packing density, and as growing or shrinking ventricular protuberances, invaginations and evaginations. Using experimental techniques in rats (short-survival, sequential survival, and long-survival ³H-thymidine autoradiography, respectively), we dated the changing proliferative dynamics of these NEP mosaics, tracked their migratory paths and settling patterns, and related that information to the time of origin of different neuronal populations in various brain regions and structures. On the assumption that the neurons that form discrete parenchymal regions and structures (different lobes, ganglia, nuclei, etc.) in the neighborhood of these NEP mosaics are progeny of those mosaics, we name the latter as putative NEPs of those structures. Examples of broader identifications of NEP “stretches” are frontal cortical NEP, hypothalamic NEP, thalamic NEP, and hippocampal NEP. An example of more specific identification of “patches” along the hippocampal NEP are subicular NEP (the mosaic that generates neurons of the subiculum), ammonic NEP (the mosaic that generates neurons the pyramidal cells of Ammon’s horn), and dentate NEP (the mosaic that generates the granule cells of the dentate gyrus). In cases where a germinal matrix abuts a fiber tract, we identify that as a putative gliopithelium (GEP), e.g., the fornical GEP. In addition to the cells that migrate short distances from their germinal sites of origin, there are others that migrate over long distances. Examples are the cells of the rostral migratory stream in the anterior telencephalon that migrate to the olfactory bulb (Altman, 1969), and the intramural (parenchymal) and extramural (subpial) migratory streams of the rhombencephalon that form the precerebellar nuclei in the medulla and pons (Altman and Bayer, 1987a-d). In such instances, we identify these NEP mosaics on the basis of the available experimental evidence of their ultimate destinations. Finally, there are many sites where we have no information at all on the destination of the neurons leaving particular NEP mosaics and indicate that uncertainty either by a question mark following a tentative identification or by omitting any identification.

Neurogenesis and its relation to the Expansion and Shrinkage of the Supraventricles. The NEP originates as an open proliferative sheet, the neural plate, without its own fluid environment. As we shall illustrate in Volume 5, the ventricles begin to form at about GW3, after the closure of the neural tube (the future spinal cord) and the cephalic vesicles (the future brain) is completed. Following that momentous morphogenetic event, the ventricles expand enormously during the rest of the first trimester and, with differences among the different ventricles, then begin to shrink during the second or third trimester, and gradually assume their mature size and configuration. The first lumen to expand is the rhombencephalic ventricle. Beginning as a shallow invagination beneath a thin membrane (the medullary velum) at about GW4, it expands greatly as a cavernous cistern is formed by the flexures of the medulla, pons and brainstem, underneath the cover of the medullary velum. As illustrated in this Volume, the expansion of the rhombencephalic ventricle continues until about GW9, and then it starts to shrink and gradually assumes the form of the familiar 4th ventricle. Next, the mesencephalic ventricle (the future aqueduct) and the diencephalic ventricle (the future 3rd ventricle) start to expand and, after a shorter developmental period, begin to shrink. Finally, at about GW5.5, the lateral ventricles begin to form as symmetrical balloon-like fluid compartments of the midline diencephalic ventricle and expand enormously up to GW11; thereafter the lumen of the lateral ventricles begin to shrink as the basal ganglia and the cerebral cortex develop. We name these greatly expanded embryonic cisterns *supraventricles* to distinguish them from the greatly diminished and transformed mature ventricles for the following reasons.

(1) Unlike the shrunken mature ventricles, the cavernous embryonic ventricles are lined by proliferating NEP cells rather than differentiated ependymal cells. This raises the possibility that the embryonic and mature ventricles and the CSF they contain serve different functions.

(2) We know since the pioneering studies of Sauer (1936) that the NEP is a pseudostratified epithelium and that the nuclei of NEP cells shuttle to the ventricular lumen before undergoing mitosis (Sauer called the process interkinetic nuclear migration). This shuttling to the lumen as a prerequisite for cell division means that the rate of NEP cell proliferation is limited by the length of the ventricular shoreline because the elongated cells straddling the depth of the pseudostratified NEP cannot divide unless there is room for them to move to the NEP/CSF interface. An interesting point in this context is the spatial orientation of the dividing cells near the lumen. It has been noted in the past (e.g., Bayer and Altman, 1991a) that the cleavage plane of these mitotic cells may be radial, or vertical (tangential to the ventricular lining) or horizontal (parallel to the ventricular lining). Vertically cleaving cells occupy double of the NEP/CSF interface when compared with hor-

izontally cleaving cells. It has been hypothesized recently that vertical cleavage results in symmetrical cell division, and horizontal cleavage in asymmetric cell division (e.g., Chenn and McConnell, 1995; Kornack and Rakic, 1995). Symmetric cell division is assumed to produce two proliferative NEP cells lying next to one another along the ventricle; collectively that should result in an expansion of the ventricular shoreline. Asymmetric cell division is assumed to produce a postmitotic daughter cell, presumably the one farther from the ventricle, a differentiating cell that is ready to leave the NEP. Some evidence has been presented recently of an increase in asymmetric division in the cortical NEP of mice as a function of increasing fetal age (Estivill-Torrus et al., 2002). Asymmetric division should not affect the length of NEP/CSF interface. The shrinkage of the NEP during fetal development would take place if more and more NEP cells lost their mitotic potency and their nuclei no longer returned to the ventricular lumen.

(3) Why is it obligatory for NEP cell nuclei to move to the ventricular lumen to divide when the stem cells of secondary germinal matrices, like the subpial external germinal layer of the cerebellum or the subgranular zone of the hippocampus, divide some distance from the ventricles? One possibility is that the embryonic CSF contains some factor or factors necessary for primary (pluripotent) NEP cell division. This is supported by reports that embryonic CSF promotes NEP cell survival, proliferation, and neurogenesis in experimental animals (Mashayekhi et al., 2002; Gato et al., 2005).

(4) The distinctiveness of the two ventricular systems is suggested by a different functional relationship between the *supraventricles* and the *fetal CP*, and the enduring *ventricles* and the *mature CP*. To begin with, the initial expansion of the rhombencephalic and the telencephalic *supraventricles* starts before the CP forms (this will be documented in Volume 5 of this series). Hence, unlike the CSF of the mature ventricular system, the CSF of the embryonic *supraventricles* must originate from some other source than CP secretion.

(5) Moreover, the fetal CP that forms later and rapidly expands to fill the rhombencephalic and telencephalic *supraventricles* (as illustrated in this Volume) has a different cellular organization than the mature CP (Kappers, 1958; Tennyson and Pappas, 1964; Shuangshoti and Netsky, 1966; Dohrmann, 1970; Dziegielewska et al., 2001; Johanson et al., 2005). The adult choroid plexus is a distinctive frond-like tissue composed of a monolayer of differentiated cuboidal cells that surround a capillary core. The exposed surface of these cuboidal cells is covered by a rich meshwork of microvilli and some cilia, and the cell interior is filled with mitochondria. These features, and added evidence, is the basis of the widely held view that the mature CP is a secretory tissue involved in CSF production. In contrast, the fetal CP is a smooth, multilayered

(pseudostratified) epithelium composed of spindle-shaped cells that have a simple exposed surface and contain few mitochondria. Unlike the mature CP cells, these fetal CP cells are filled with glycogen. Hence, it has been suggested that the principal function of the embryonic CP is the glycolytic (anaerobic) support of NEP cell proliferation and neurogenesis rather than the production of CSF.

(6) Finally, and most importantly, the chronological differences in the expansion, configurational transformations, and persistence of the different superventricles can be related to regional differences in the kinetics, date of origin, and time span of cell proliferation in stretches and patches of the NEP lining that produce neurons and neuroglia for different brain structures. For instance, the dorsal roof and lateral wall of the telencephalic superventricle is formed by an extensive, continuous dome-like stretch of NEP that generates an immense number and homogeneous set of cortical neurons in a precise sequential order (infragranular layer cells first, granular layer cells next, and supragranular layer cells last) through its entire extent and over an extended period of time. These neurogenetic features account for the smooth, spherical configuration of the NEP lining of the telencephalic superventricle over an extended period during embryonic and fetal development. In contrast, the ventral and medial shores of the telencephalic superventricles are much less regular because discrete NEP stretches or patches generate neurons here for diverse brain structures – septum, basal ganglia, nucleus accumbens, basal telencephalon, amygdala, hippocampus, etc. – each with different neurogenetic timetables, cell compositions, and population sizes. At these sites, the expanding and shrinking NEP mosaics (and the differentiating parenchymal structures associated with them) produce variably shaped ventricular pools, recesses, and narrows. Still more complex are the configurational changes over time in the diencephalic, mesencephalic, and rhombencephalic superventricles. This is so because at these locations short NEP patches give rise to a multitude of structurally and functionally different brain nuclei, many of which are composed of a relatively small number of neurons generated over a short time span. For instance, we can distinguish along the diencephalic superventricle not only NEP stretches that give rise to the thalamus, epithalamus, subthalamus and hypothalamus, but also patches – temporary evaginations and invaginations – that generate neurons, for instance, for the distinct thalamic nuclei. These expanded and short-lived shorelines create an expanded NEP/CSF interface for the optimal generation of neural stem cells of a particular type.

Glioepithelia and the Ependymal Linings of the Enduring Ventricles. Glioepithelia (GEP) are fate-restricted tissues of proliferative cells that produce neuroglia. On the basis of experimental studies in rats, in which proliferative cells are tagged with ^3H -thymidine, we distinguish four types of GEP. The first is difficult to distinguish from the

primary NEP lining the ventricles except that these patches tend to be thin and display proliferative activity for some time after local neurogenesis has ended. These patches begin to appear caudally in the older fetuses of the age group covered in this Volume, and because at most of these sites an ependymal lining will be forming around the enduring ventricles, they are designated uncertainly as glioepithelium/ependyma (G/EP). Examples are the medullary, pontine, hypothalamic, and thalamic G/EP. The second type lines fiber tracts that are devoid of neuronal cell bodies. In the older fetuses of this age group this type is exemplified by the fornical GEP, a continuation of the hippocampal NEP. The fornical GEP probably contains precursors of oligodendrocytes. The third type is the perifascicular GEP that surrounds and penetrates large fiber tracts. In the older specimens of this age group these are seen around the olfactory tract and the optic nerve, chiasm and tract. The fourth type, the subpial GEP, is found as a covering of the telencephalon, known as the subpial granular layer (Brun, 1965). The perifascicular and subpial GEPs may be continuous and of placodal rather than neuroepithelial origin.

Secondary Germinal Matrices, Migratory Streams, and Sojourn Zones. Prominent transitional structures of the developing brain during the second trimester, as illustrated in Volume 3 of this series, are the secondary germinal matrices, the prominent sojourn zones, and the large migratory streams. The principal secondary germinal matrices are the subventricular zone (SVZ) abutting the cortical NEP, the SVZ of the basal ganglionic (anterolateral, anteromedial and posterior) eminences, the subgranular zone of the hippocampal dentate gyrus, and the external germinal layer of the cerebellar cortex. All of these secondary matrices are late-generated, fate-restricted stem cells that divide some distance from the ventricles and produce microneurons (neurons with small cell bodies and locally ramifying axons); many of which are known as “granule cells.” In the specimens we use, the time of emergence of the cortical SVZ could not be determined because of the difficulty of distinguishing it from the NEP, but it is clearly present by GW11. The basal ganglionic SVZ begins to form as early as GW7.5 and is prominent between the NEP and the parenchyma by GW8. The external germinal layer that originates in the germinal trigone of the cerebellar NEP is not recognizable until GW9. The hippocampal dentate NEP, which is the source of the dentate migration (and which will later form the subgranular zone) is present by GW9; however, the dentate gyrus is only a miniscule structure at GW11. In general, there is no evidence for the onset of microneuron production during this period.

There are long and short migratory streams that carry neural stem cells and/or immature neurons from the NEP to their destination. A prominent long-distance migration during the second trimester is the rostral migratory stream of the telencephalon which, among others, contains cells

that differentiate as olfactory bulb granule cells. The rostral migratory stream is not clearly distinguishable at GW9 from the olfactory NEP, which generates the output neurons (mitral cells) of the olfactory bulb; it becomes recognizable as a distinct entity by GW11, and expands greatly during the second trimester (Volume 3). Among the long-distance precerebellar migratory streams, the posterior intramural migratory stream, which contains the neurons that migrate to form the prominent inferior olive, is evident by GW7.5, and remains so throughout the late first trimester. Also present in these specimens are the posterior extramural migratory stream that contains neurons of the lateral reticular and external cuneate nuclei, and the anterior extramural migratory stream that contains pontine gray neurons. However, the pontine gray nucleus is not evident until GW11 when the earliest corticofugal fibers begin to traverse it and the earliest fibers of the middle cerebellar peduncle begin to form.

In Volume 3 during the second trimester, we have illustrated the prominence of the cortical stratified transitional field (STF) sandwiched between the NEP and the cortical plate, the future gray matter. We have identified six distinct cellular and fibrous layers within the STF, where cortical neurons sojourn for some time and mingle with afferent, efferent, and commissural fibers before they resume their migration and settle in the cortical plate. We postulated that the STF is a staging area where connections are formed between unspecified cortical neurons and the functionally and topographically specified thalamocortical fiber systems that provide input to them. As seen in the present Volume, the cortical STF is developing during the late first trimester. STF1 and STF5 are first evident at GW8 in the earlier-maturing anterolateral cortical region spreading into the later-maturing dorsomedial cortical region. The formation of STF5 appears to coincide with the growth of thalamocortical fibers through the diencephalic-telencephalic junction and their penetration into the formative cerebral cortex. Between GW9 and GW11 an additional layer, STF4, begins to emerge slowly and uncertainly in the earlier maturing regions of the cortex (i.e., the lateral aspect of the anterior hemisphere). The emergence of STF4 may be associated with the onset of the descent of corticofugal fibers. The other STF layers (STF3, STF2, and STF6) do not start to form until the beginning of the second trimester (Volume 3).

Several other transitional fields are seen in the developing brain during this period. In the GW7.5-GW8 specimens, large cell aggregates surround the thalamic NEP that we interpret to be sojourning neuronal populations of the anterior, dorsal and ventral thalamic complexes. And, based on observations in rats, we assume that the posterior thalamic complex that will produce neurons for the lateral geniculate body and the pulvinar is initially in a dorsomedial position and then migrates gradually ventrolaterally toward the growing optic tract. The morphogenetic sig-

nificance of the transitional cell columns and fiber bands seen in the GW9 thalamus, before the thalamic neurons assume their “nuclear” configuration, remains to be elucidated. Another region with alternating bands of sojourning cells and growing fibers is the cerebellar transitional field (CTF). Six layers are distinguished in the formative cerebellum in GW7.5 specimens from the surface toward the NEP: the fibrous CTF1, the cellular CTF2, CTF3 with cells and fibers, the cellular CTF4, CTF5 with cells and fibers, and the cellular CTF6. On the basis of experimental evidence in rats, we interpret the upper cellular layers (CTF2-4) as the earlier generated deep neurons that sojourn for a while superficially, and the lower cellular layers (CTF5-6) as consisting mainly of the later generated Purkinje cells. This stratification becomes blurred by GW8 and GW9 as the Purkinje cells migrate toward the surface and the deep neurons move back toward the core of the cerebellum. The upward migration of Purkinje cells is associated with the spreading of the external germinal layer over the surface of the formative cerebellar cortex. By GW11 most of the Purkinje cells are in a superficial position, where they form parasagittal bands beneath the continuous canopy of the external germinal layer.

F. A Note on Genetic Analyses of NEP Mosaicism

As we noted earlier, the NEP cells that line the banks of the supraventricles look alike in Nissl-stained preparations. The idea of NEP mosaicism is based on the site- and age-dependent distinctiveness of stretches and patches of the NEP in terms of their thickness, the temporary formation of larger protuberances and smaller evaginations and invaginations into the ventricle, and their differential proliferative dynamics, as ascertained with ³H-thymidine autoradiography. These spatiotemporal differences in the regional appearance and proliferative dynamics of NEP mosaics, and the fact that they give rise to different brain structures and cell types, raise the possibility that the stem cells composing them are different genetically before they start to differentiate. There is, indeed, emerging experimental evidence in embryonic and fetal mammals (mostly mice) for the genetic distinctiveness of some of the stretches and patches of NEP that are homologous with those we describe here in the first trimester human brain. The experiments are based mostly on *in situ* hybridization of homeodomain-containing transcription factors that are visualized in the developing brain of normal and mutant mice with immunohistochemical markers. Several laboratories, for instance, have reported a pronounced expression of *Pax6*, in combination with other transcription factors, in the dorsal neocortical NEP (e.g., Walther and Gruss, 1991; Stoykova and Gruss, 1994; Warren et al., 1999; Estivill-Torrus et al., 2002; Kimura et al., 2005). At the embryonic age when *Pax6* expression is limited to the dorsal telencephalon, *Nkx2.1* and *Gsh2* are expressed in the ventral telencephalon and the medial telencephalon (Corbin et al.

2003). Some differences were noted in the expression of these factors in the lateral, medial and posterior components of the ganglionic ventral telencephalon. According to another study, different markers highlight different components of the medial telencephalon and the diencephalon in 12.5 day-old mouse embryos (Kimura et al., 2005). Using a different terminology from that used by the authors, *Wnt8b* appears to demarcate the entire hippocampal primordium in the medial cortex, and the epithalamus and the mammillary body in the diencephalon. Within the domain of the medial telencephalon, *Ephb1* demarcates the Ammonic NEP, *Wnt3a* the dentate NEP, and *TTR* the primordium of the telencephalic choroid plexus. Although as yet this genetic approach is limited to a single species in mammals, it is expected that it will shed considerable light on NEP cell heterogeneity and the regional differentiation of the embryonic and fetal brain in other animals and man.

G. A Note on Functional Maturation

There is currently considerable scientific interest in the physiological maturation of the prenatal human brain, and the correlated issue of the mental status of the embryo and fetus is receiving much public attention. Studies in the first half of the twentieth century with aborted fetuses have indicated that embryos of about 20-21 mm CR length (corresponding to the GW7.5 specimens in the present volume) begin to reliably respond to tactile stimulation with *holokinetic* (“total pattern”) body movements (Fitzgerald and Windle, 1942; Hooker, 1942). In GW10 fetuses (CR 48.5 mm) *ideokinetic* or isolated movements are also elicited, such as partial closure of the fingers (though not effective grasping), when the palm of the hand was stimulated (Humphrey, 1964). The more recent introduction of ultrasonic recording techniques has permitted the observation of the emergence of “spontaneous” fetal behavior in normal embryos and fetuses *in utero*. A pioneering study showed that the holokinetic “startle” response emerges as early as GW6, isolated arm and leg movements by GW7, and head rotation and hand and face contact beginning at about GW8 (de Vries, 1982, 1985). According to a more recent study with improved ultrasonic recording methods, isolated arm movements are more frequent than isolated leg movements during the first trimester, and head turning and hand to head contact do not occur with high frequency until the second trimester (Kurjak et al., 2005).

Are the late-embryonic and early-fetal movements reflex reactions mediated by lower-level spinal cord and brain stem mechanisms, or are they emerging voluntary activities carried out under higher-level cortical guidance, reflecting sentience? We have raised this question earlier in the context of our study of the development of the reflex circuitry of the human spinal cord (Altman and Bayer, 2001). Because the spinal cord substrate of the sensorimotor reflex arc begins to form between GW7 and GW8 – the collateral branches of dorsal root sensory nerves sprout and then

reach the ventral horn motor neurons during this period – we proposed that the isolated limb movements displayed by embryos of that age are reflex reactions. The morphogenetic evidence presented in this Volume extends that inference by showing that the first trimester embryonic movements cannot be cortically mediated voluntary activities. First, the higher-level sensory channel, the medial lemniscal system relayed in the thalamus, does not reach the cortical plate (the future cortical gray matter) during this period. The internal capsule that contains the thalamocortical fibers is not yet evident at GW7.5 either in the lateral border of the thalamus or the narrow parenchymal bridge that links the diencephalon and the telencephalon. A large collection of thalamocortical fibers “funnel” through the internal capsule by GW8, passing through the basal ganglia and approaching the base of the formative cerebral cortex. These fibers begin to penetrate STF4 of the paracentral lobule by GW9 and that process continues through GW11. It is probable, but this needs to be experimentally verified, that the neurons of layer IV of the cortex, the principal target of thalamocortical fibers, are either still being generated or are still in the STF5 sojourn zone. Thus, there is as yet no functional connection between the thalamus and cells of the cortical plate. The second consideration concerns corticofugal output to the brain stem and spinal cord, which is a prerequisite for cortically mediated motor control. The pontine gray, through which the corticospinal tract descends to the spinal cord, is not evident at GW9; it begins to form about GW11. In one such specimen illustrated (Y1-59; CR 60 mm), but not in the other (C1500; CR 57 mm), a bundle of corticofugal fibers traverses the pons. Although the earliest contact between thalamocortical fibers and differentiating cortical neurons may be established during this period, we have shown earlier that the lateral and ventral corticospinal tracts do not penetrate the spinal cord until the second trimester (Altman and Bayer, 2001: Figs. 7-23, 7-24, 7-35) and are still unmyelinated at birth (Altman and Bayer, 2001: Fig. 8-14).

H. References

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- Walther, C. and P. Gruss (1991) *Pax-6*, a murine paired box gene, is expressed in the developing CNS. *Development*, 113:1435-1449.
- Warren, N., D. Caric, T. Pratt, J. A. Clausen, P. Asavaritikrai, J. O. Mason, R. E. Hill and D. J. Price (1999) The transcription factor, *Pax6*, is required for cell proliferation in the developing cerebral cortex. *Cerebral Cortex*, 9:627-635.

PART II: GW11 CORONAL

This specimen is a stillborn female fetus with a crown-rump length (CR) of 60 mm estimated to be at gestational week (GW) 11 (Yakovlev case number RPSL-WX-1-59, referred to here as Y1-59). The brain was cut in the coronal (frontal) plane in 35- μ m thick sections and is classified as a Normative Control in the Yakovlev Collection (Haleem, 1990). Since there is no photograph of this brain before it was embedded and cut, a specimen from Hochstetter (1919) that is comparable in age to Y1-59 is used to show the approximate section plane and external features of a GW11 brain (**Figure 1**). Photographs of 22 Nissl-stained sections (**Levels 1-22**) are shown at low magnification in **Plates 1-21**. **Plates 22-35** show high-magnification views of various parts of the brain from the cerebral cortex (**Plates 22-23**) to the midbrain, pons, and medulla (**Plates 33-35**). Several high-magnification plates are rotated 90° (landscape orientation) to more efficiently use page space. Y1-59 has more mature brain structures in the diencephalon, midbrain, pons, and medulla than any other specimen in this Volume. Immature brain structures predominate in the telencephalon and the cerebellum.

Throughout the cerebral cortex, the *neuroepithelium* and *subventricular zone* are prominent. The *stratified transitional field (STF)* contains mainly *STF1* and *STF5* throughout; with *STF4* only in lateral areas and a questionable *STF2* in a few areas; *STF6* and *STF3* are not present. The *STF* is filled with migrating and sojourning neurons and, unlike any specimen in Volume 3, *has no regional heterogeneity*. The cerebral cortex is completely smooth except for a questionable calcarine sulcus in the left cerebral hemisphere (**Plate 15**); “lobes” are identified as future lobes. The most prominent developmental feature of the cerebral cortex is that both the *STF* layers and the cortical plate have a pronounced lateral (thicker) to medial (thinner) maturation gradient. There is no corpus callosum. The olfactory bulb is beneath the anterior septum and striatum; it contains the *rostral migratory stream* in its core. In anterolateral parts of the cerebral cortex, streams of neurons and glia appear to leave *STF4* and enter the *lateral migratory stream*. The hippocampus is in an immature position dorsal to the thalamus and medial to the temporal lobe. Cells are entering Ammon’s horn pyramidal layer in the *ammonic migration*, and granule cells and their precursors are migrating to the hilus of the presumptive dentate gyrus in the *dentate migration*; there is no granular layer. A massive *neuroepithelium/subventricular zone* overlies the amygdala, nucleus accumbens, and striatum (caudate

and putamen) where neurons (and glia) are being generated. The caudate, the putamen, and basolateral parts of the amygdala are smaller than in the GW13.5 specimen in Volume 3, but are similar to that specimen because the *stratial neuroepithelium* and *subventricular zone* have indistinct subdivisions. The *strionuclear glioepithelium* forms definite continuities with the fornical glioepithelium in the telencephalon. Unlike the GW13.5 specimen, the septum in Y1-59 has a *neuroepithelium* at the ventricular surface instead of a *glioepithelium/ependyma*.

The cerebellum is a thick, smooth plate overlying the posterior pons and medulla. However, there is only a thin *glioepithelium/ependyma* at the ventricular surface, indicating that all deep neurons and Purkinje cells have been generated. The deep neurons are in place beneath the cortex, but have indefinite nuclear subdivisions. The cortical surface is covered by an *external germinal layer (egl)* that is actively producing neuronal stem cells, granule, stellate, and basket cells of the cerebellar cortex. Lamination in the cortex is nearly absent, except for a thin molecular layer beneath the *egl*. Nearly all Purkinje cells are migrating, some in discrete clumps. Lobulation has barely begun in the vermis and is nearly absent in the hemispheres. The *germinal trigone* is prominent in the dorsal rhombic lip.

The third ventricle, aqueduct, and fourth ventricle are lined by a thin *glioepithelium/ependyma* indicating that neurogenesis in the primary neuroepithelium is complete. In the medulla, there are two active germinal sites in anterior and posterior parts of the ventral rhombic lip. 1) The *auditory neuroepithelium* generates cochlear nucleus neurons. 2) A large *precerebellar neuroepithelium* generates precerebellar (mainly pontine gray) neurons.

Neurons throughout the diencephalon are settling in fairly well-defined nuclear divisions; the major exceptions are the immature appearance of the lateral and medial geniculate bodies in the posterior thalamus and the hypothalamic medial mammillary body. Neurons are settled in the midbrain tegmentum, pons, and medulla. But the pontine gray is nearly absent, and neurons are still migrating into it from the large *anterior extramural migratory stream*. *Posterior extramural and intramural migratory streams* contain lateral reticular neurons, external cuneate nucleus neurons, and inferior olive neurons. The corticospinal tract forms a small cerebral peduncle in the midbrain, but has not grown into the pons.

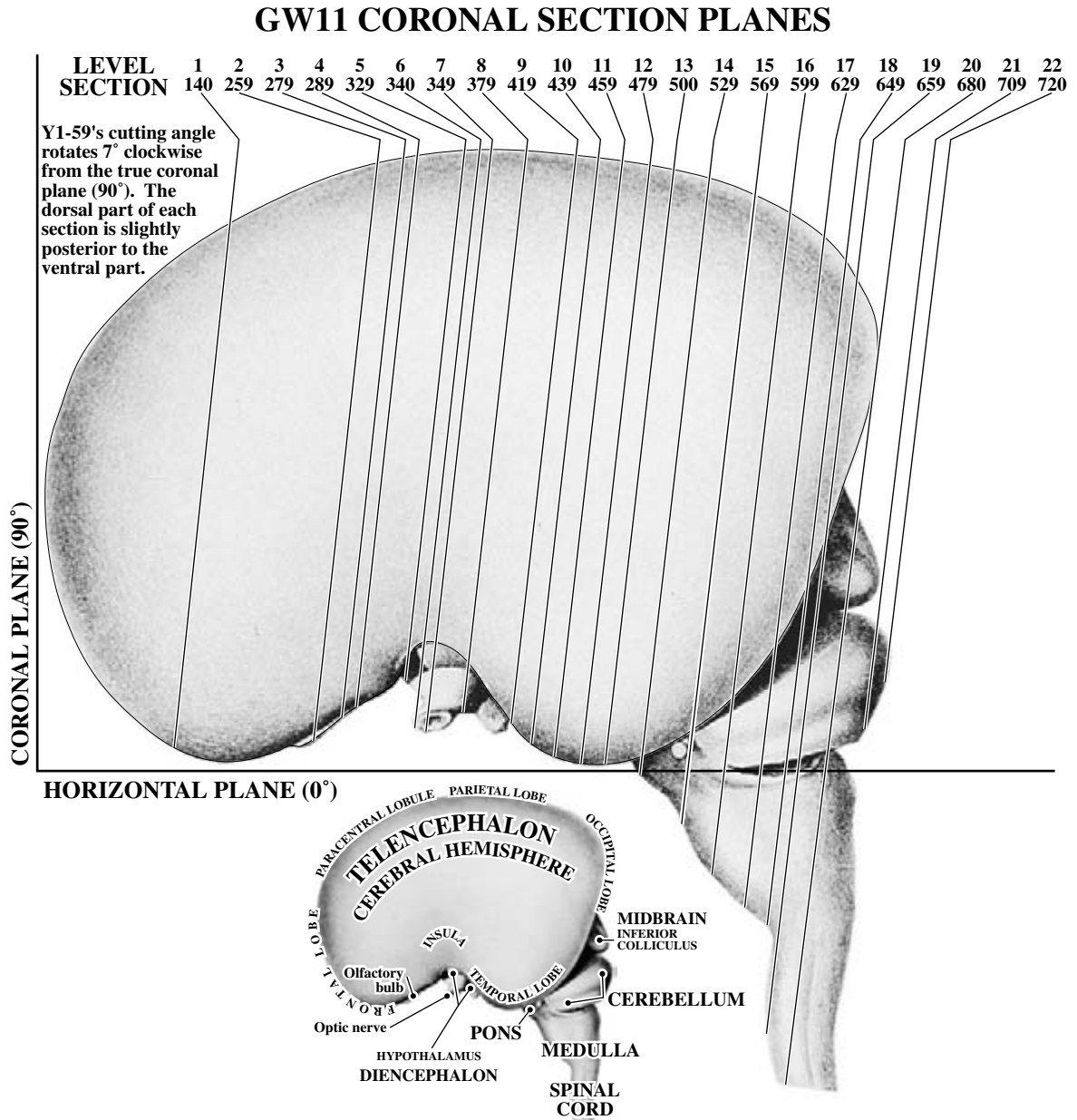


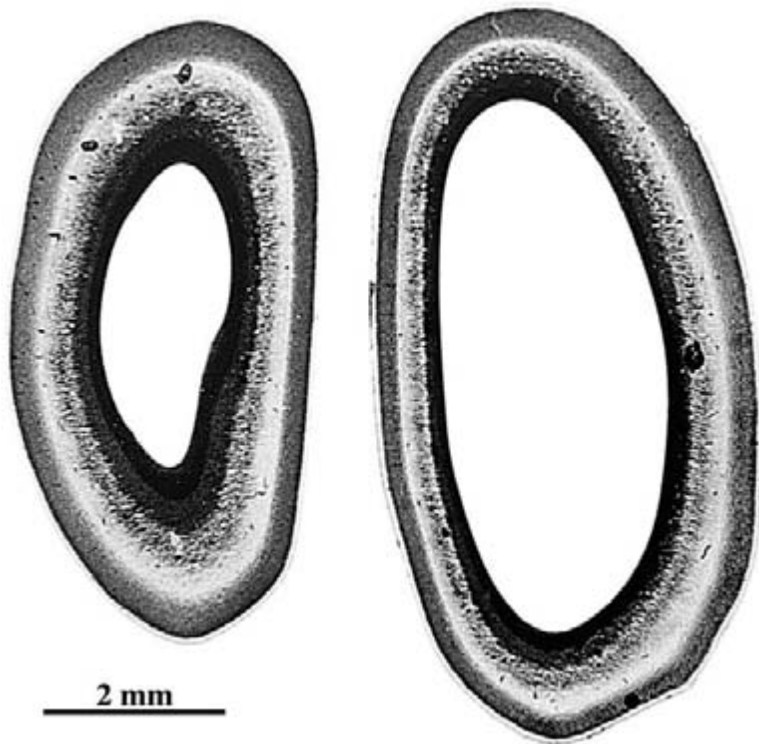
Figure 1. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 68 mm (modified from Figure 47, Table VIII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of Y1-59 in the following pages. The small inset identifies the major structural features. The cut beneath the cerebellum is the edge of the medullary velum.

PLATE 1A

**GW11 Coronal
CR 60 mm, Y1-59
Level 1: Section 140**

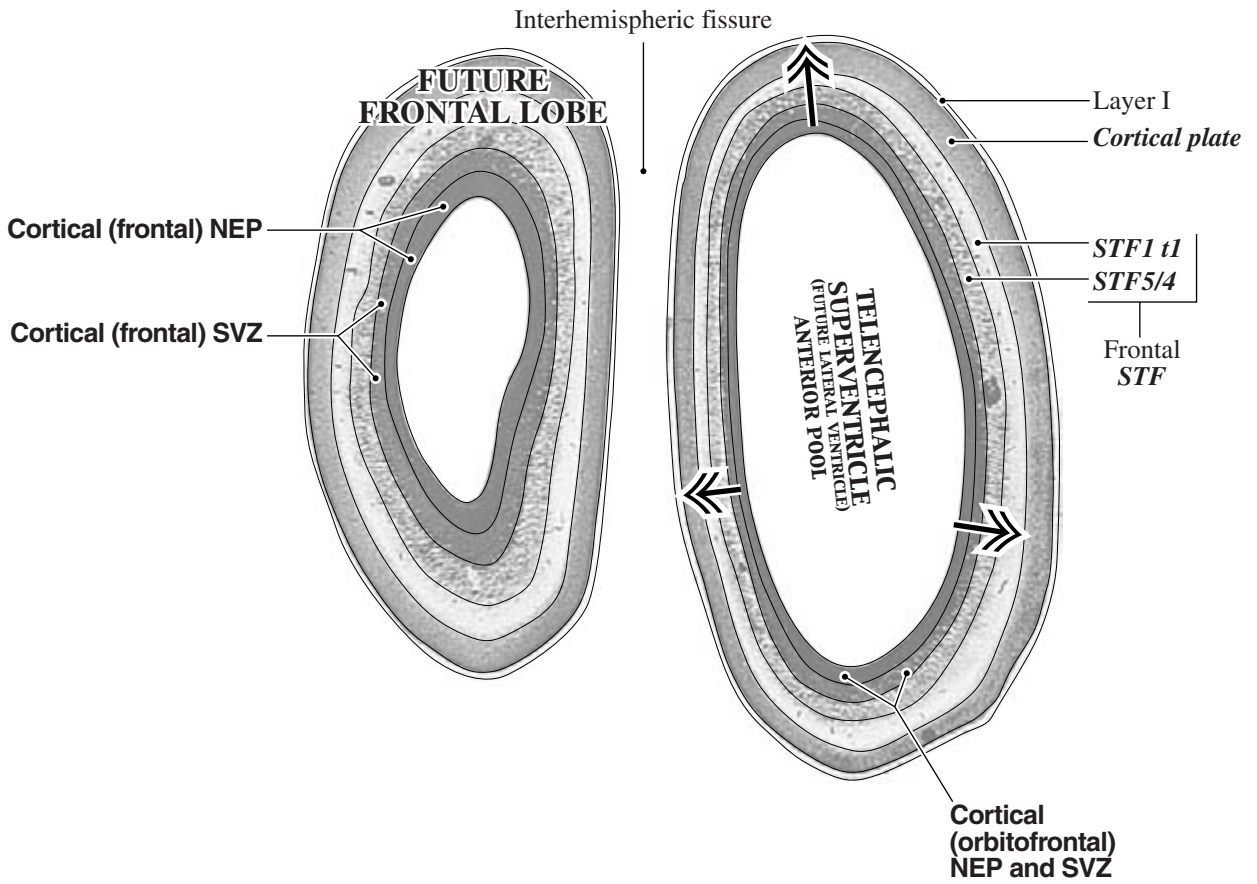
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>11</i>) when many cells are migrating through it, followed by a late stage (<i>12</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
 VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
 NEP - Neuroepithelium
 SVZ - Subventricular zone



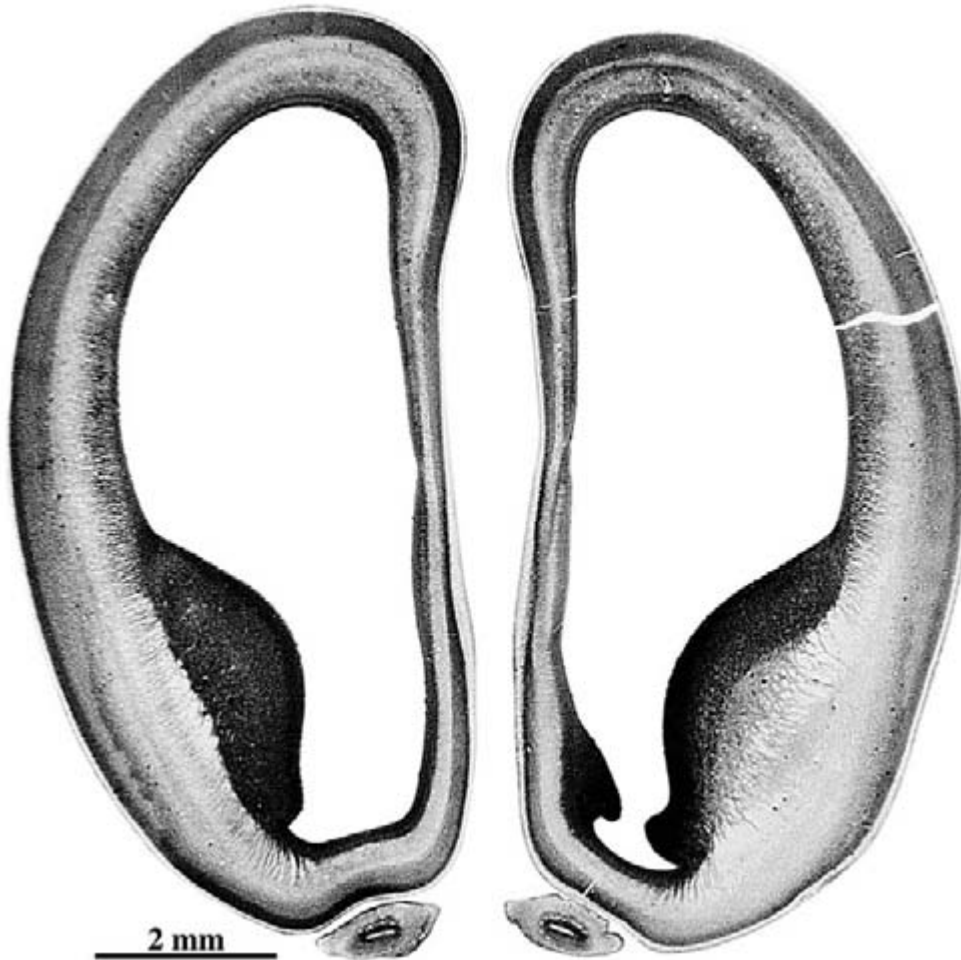
Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 2A

**GW11 Coronal
CR 60 mm, Y1-59
Level 2: Section 259**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

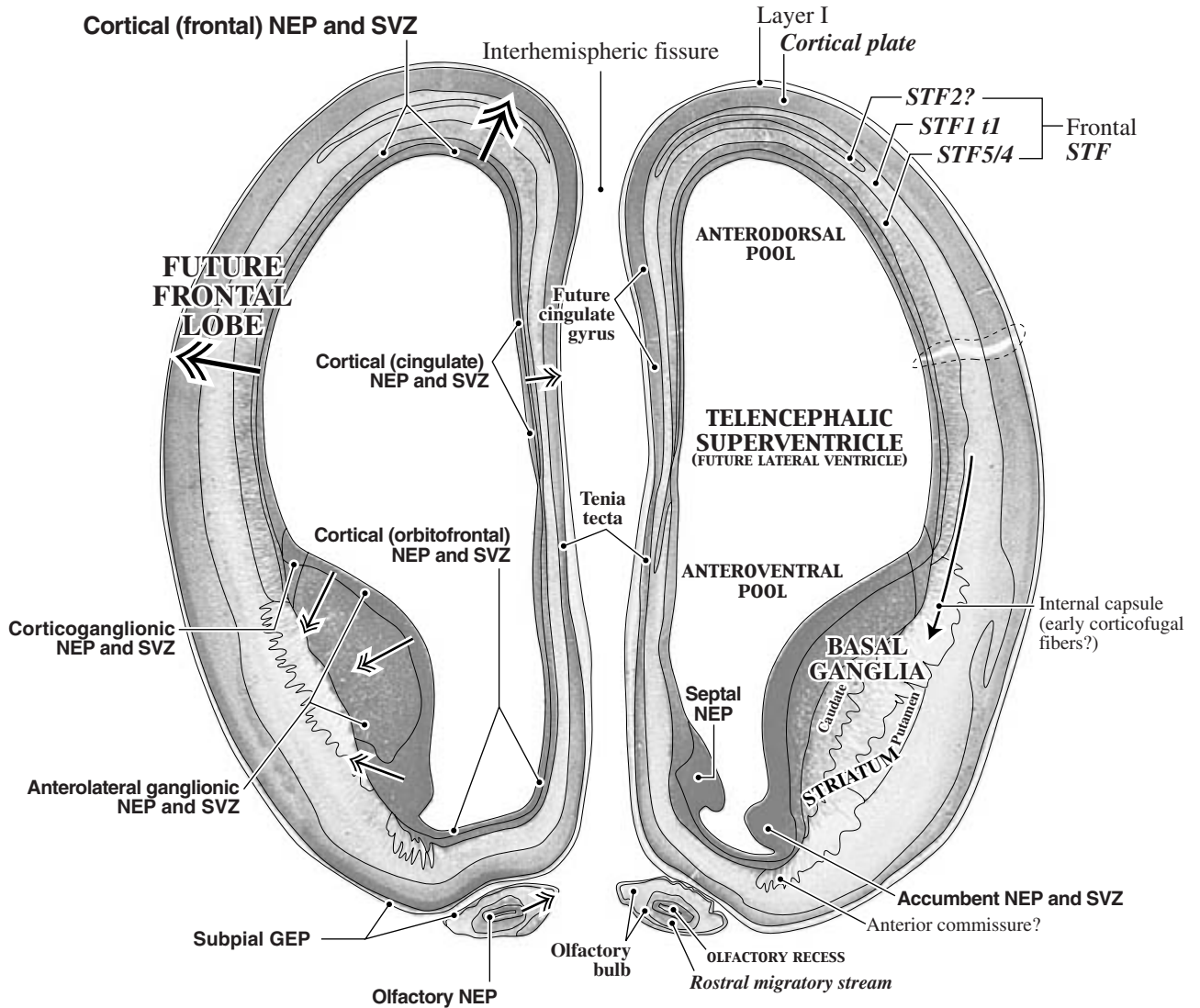
STF1 Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2 Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3 Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6 Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the frontal cortex from section 269 in Plates 22A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

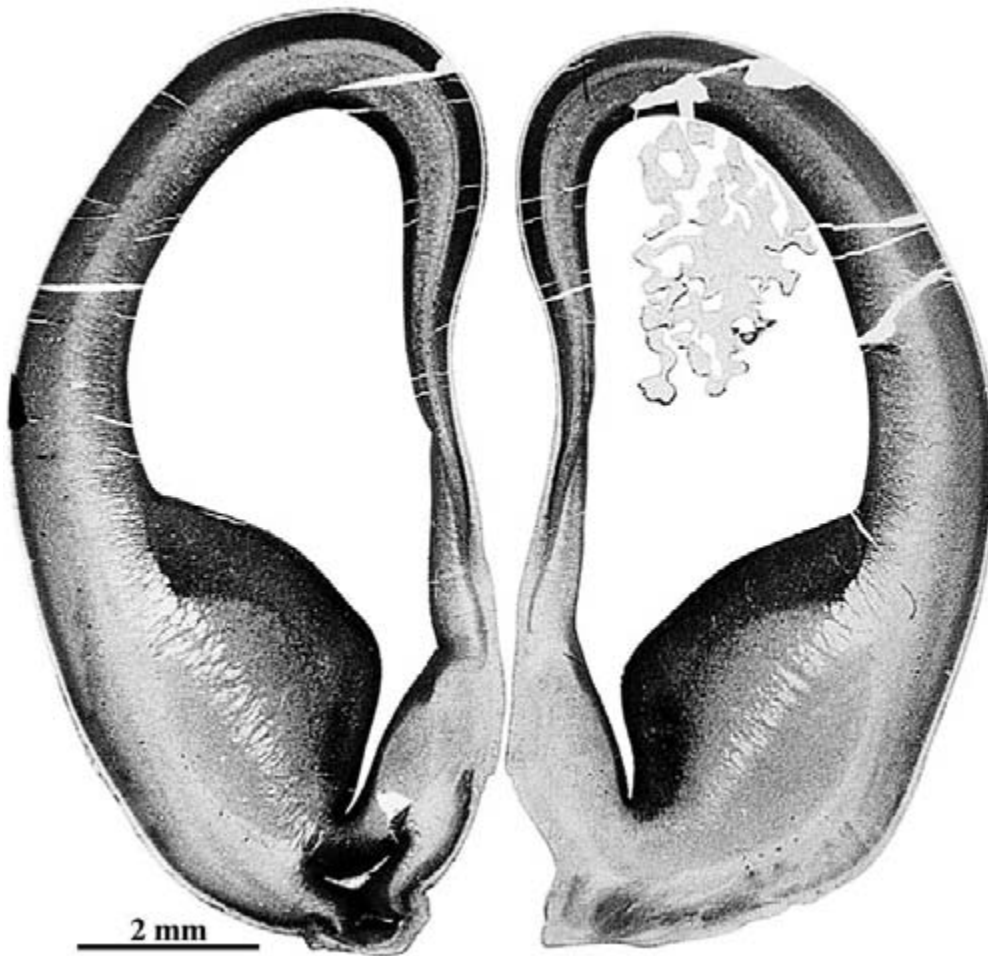
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 3A

**GW11 Coronal
CR 60 mm, Y1-59
Level 3: Section 279**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

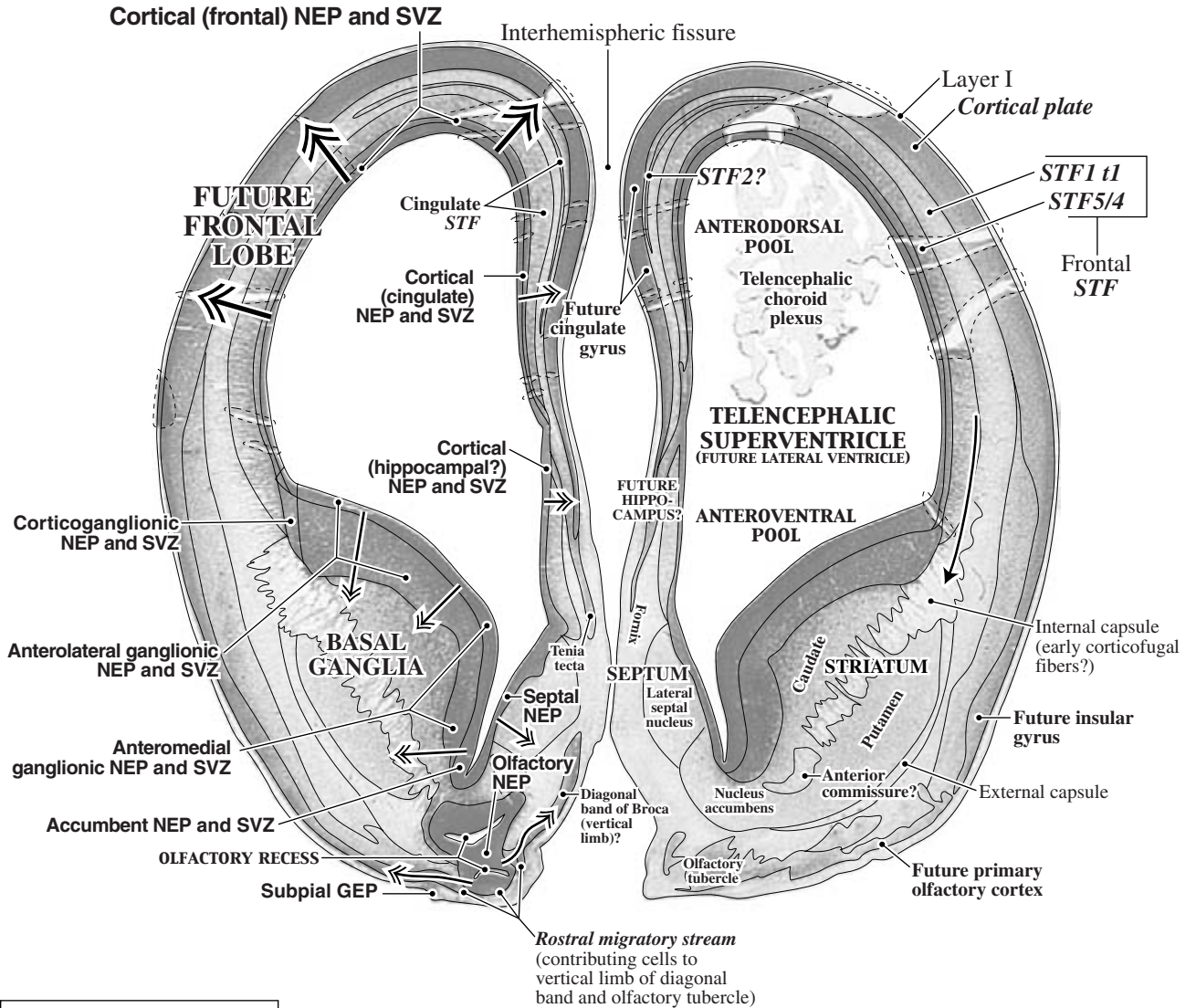
STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the frontal cortex from section 269 in Plates 22A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

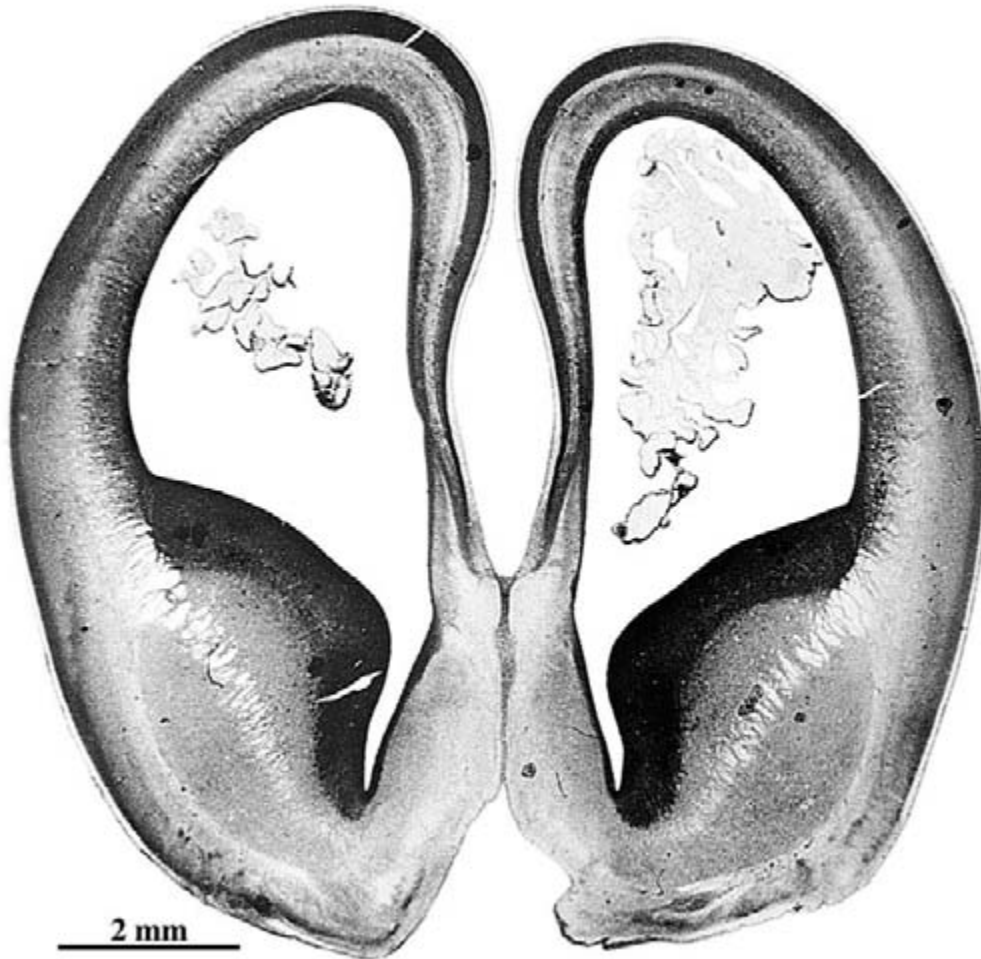
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 4A

**GW11 Coronal
CR 60 mm, Y1-59
Level 4: Section 289**

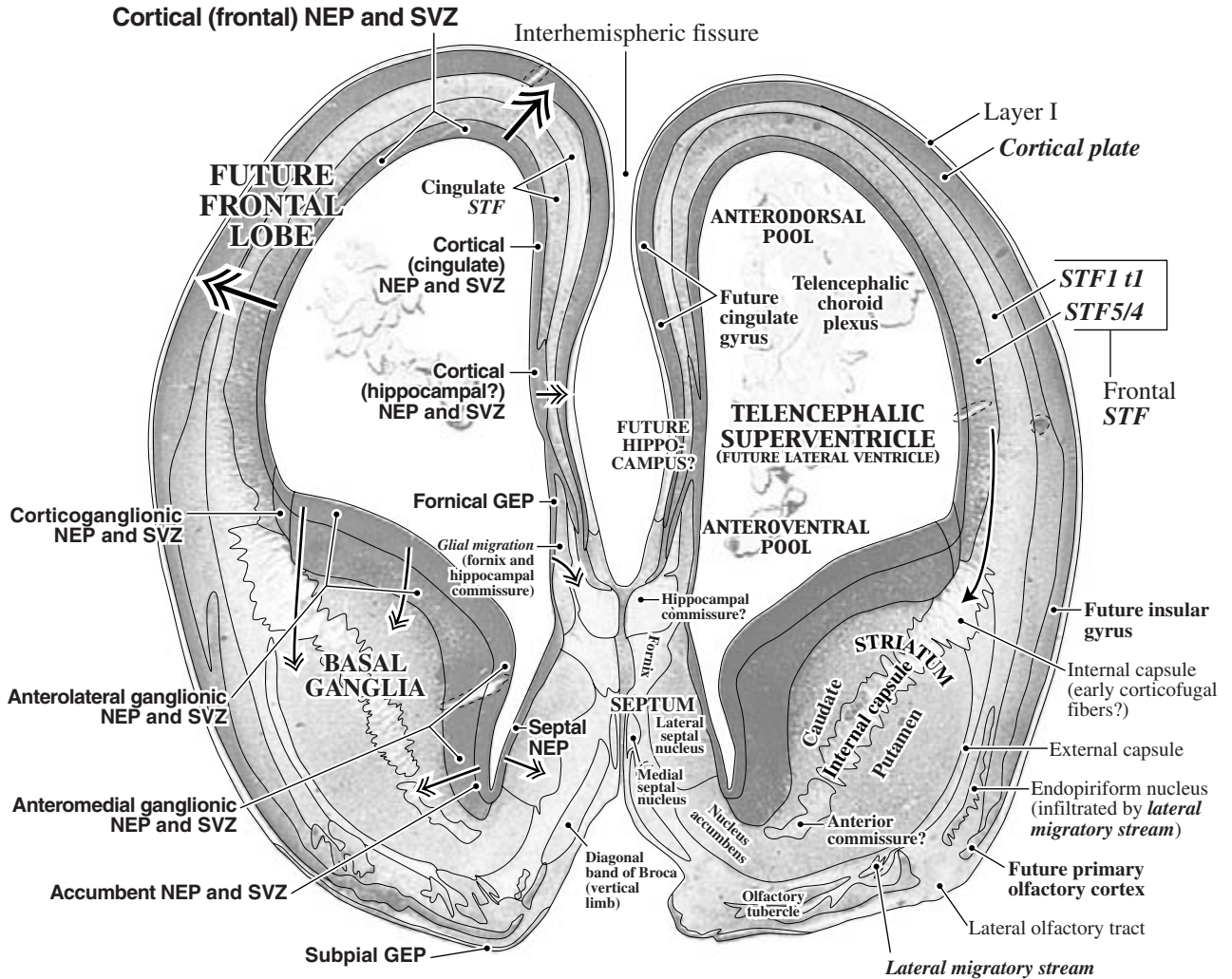
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

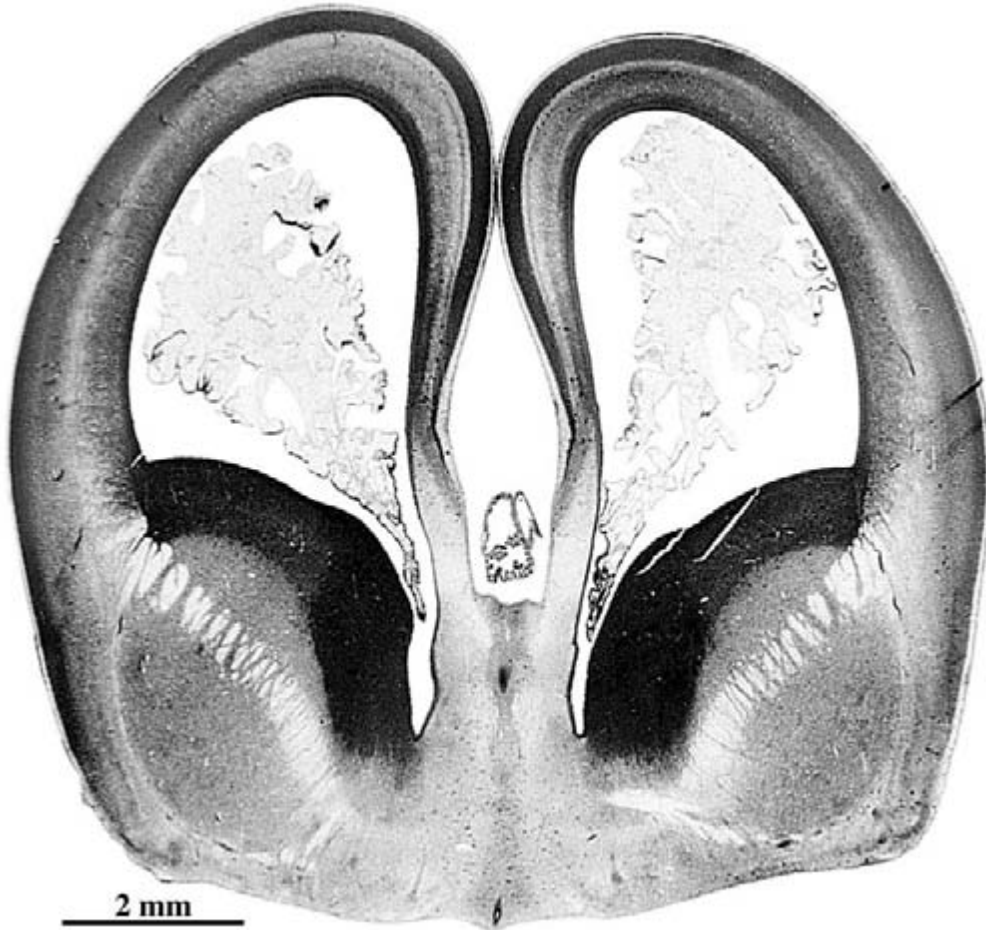
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 5A

**GW11 Coronal
CR 60 mm, Y1-59
Level 5: Section 329**

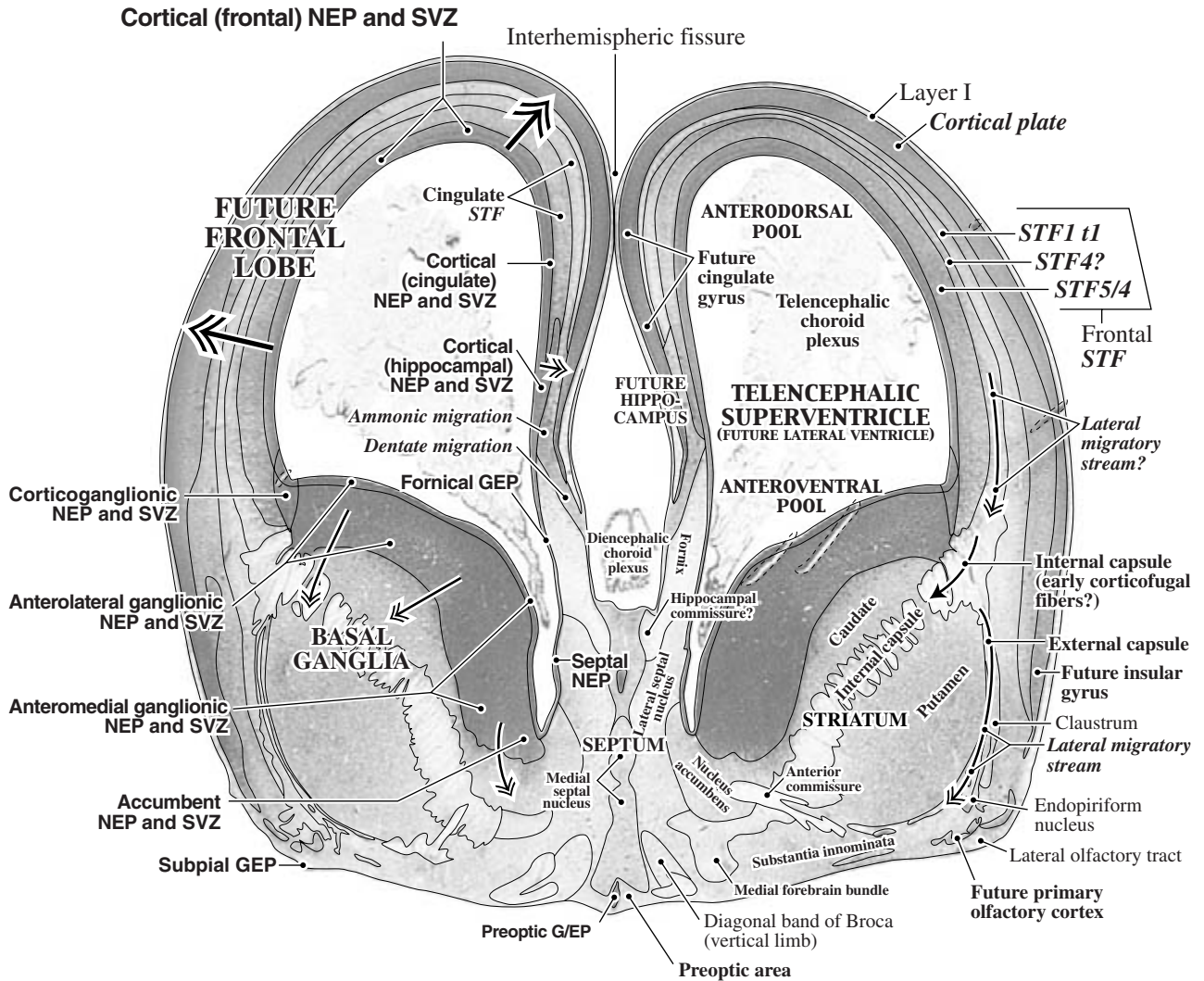
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

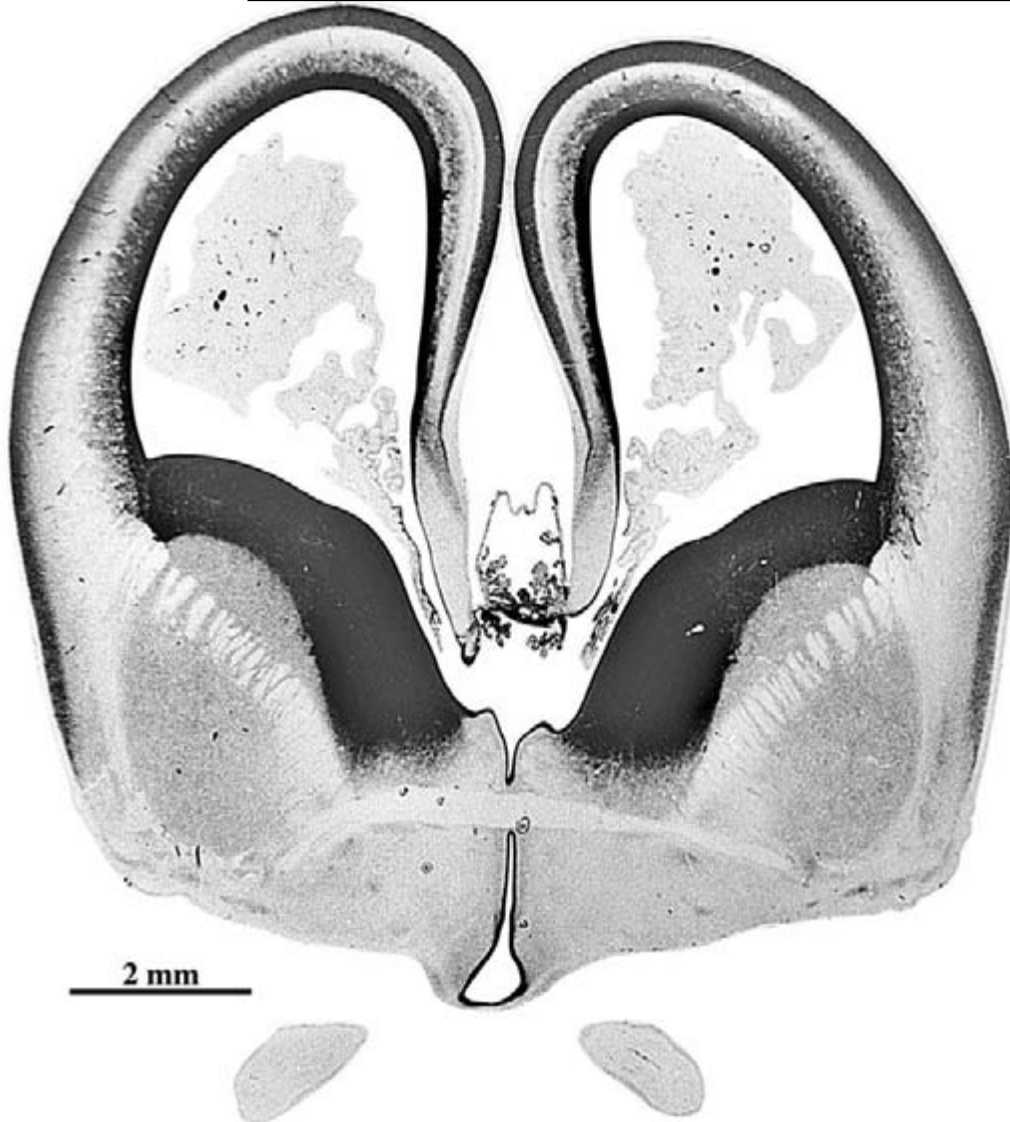
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 6A

**GW11 Coronal
CR 60 mm, Y1-59
Level 6: Section 340**

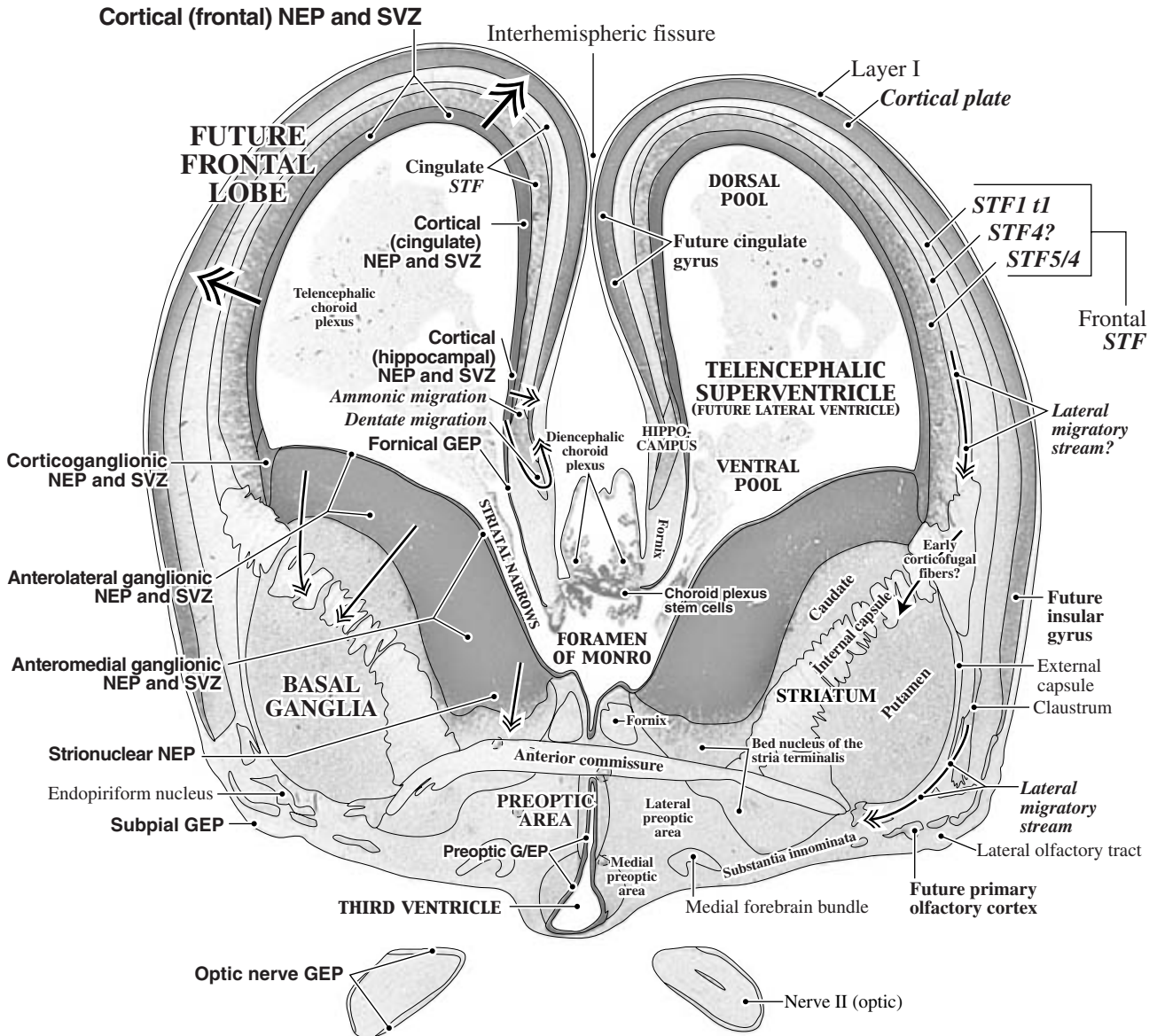
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- | | | | |
|-------------|--|-------------|---|
| STF1 | Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter. | STF4 | Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons. |
| STF2 | Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate. | STF5 | Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix. |
| STF3 | Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices. | STF6 | Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester. |



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - *Helvetica bold*
 Transient structure - *Times bold italic*
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

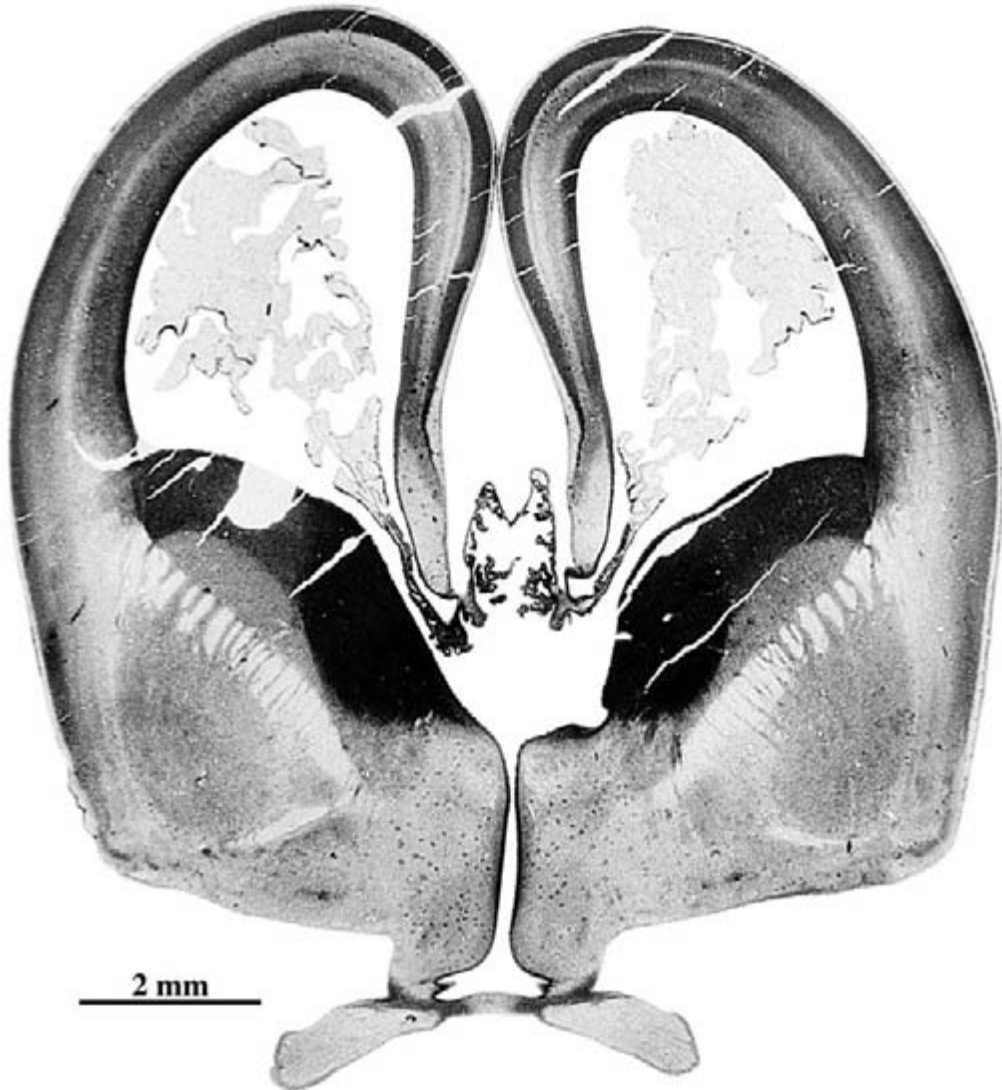
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 7A

**GW11 Coronal
CR 60 mm, Y1-59
Level 7: Section 349**

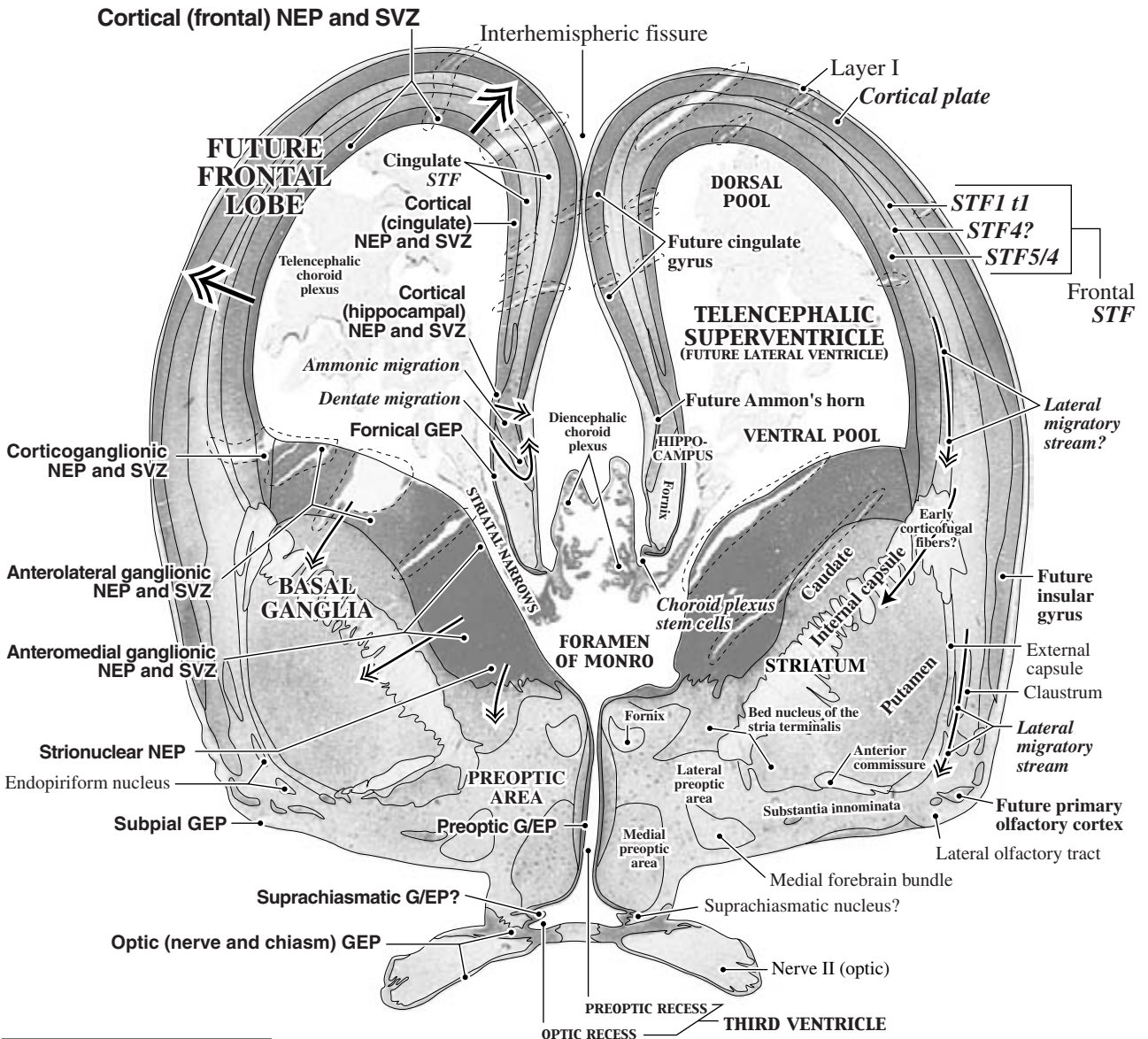
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - **Helvetica bold**
 Transient structure - *Times bold italic*
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

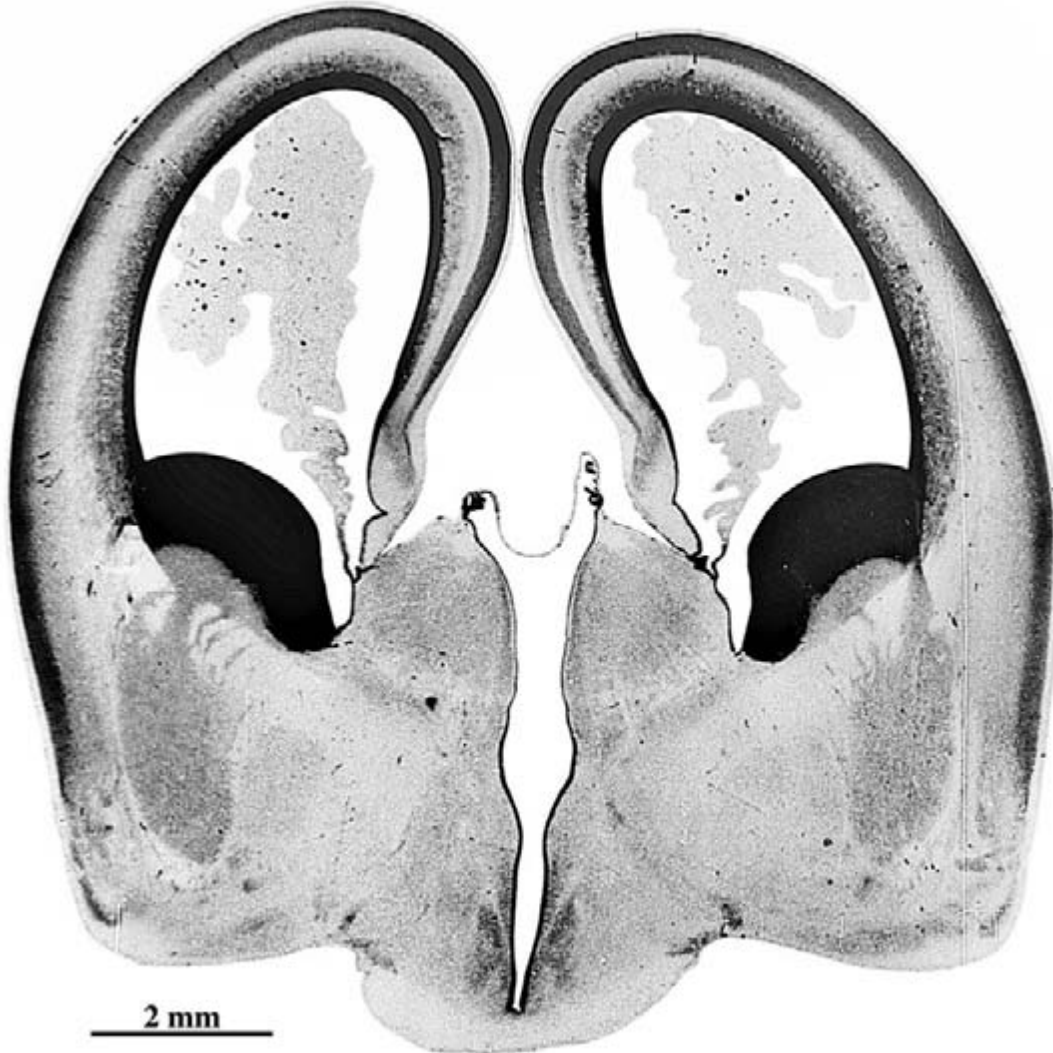
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 8A

**GW11 Coronal
CR 60 mm, Y1-59
Level 8: Section 379**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.

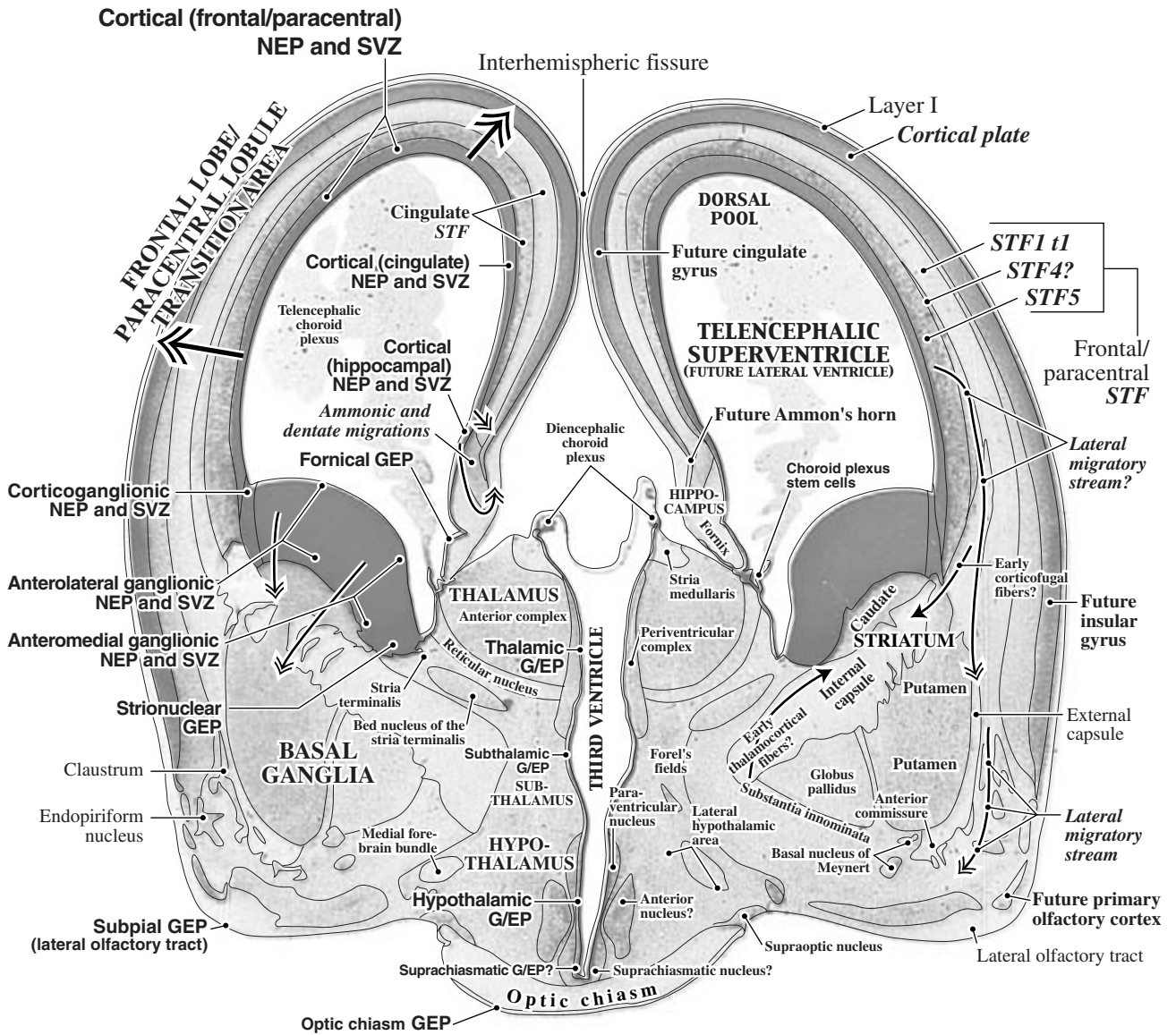


See high-magnification views of the right paracentral cortex, thalamus, and basal ganglia from section 399 in Plates 24A and B to 25A and B.

See a high-magnification view of the diencephalon and basal telencephalon from section 389 in Plates 26A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

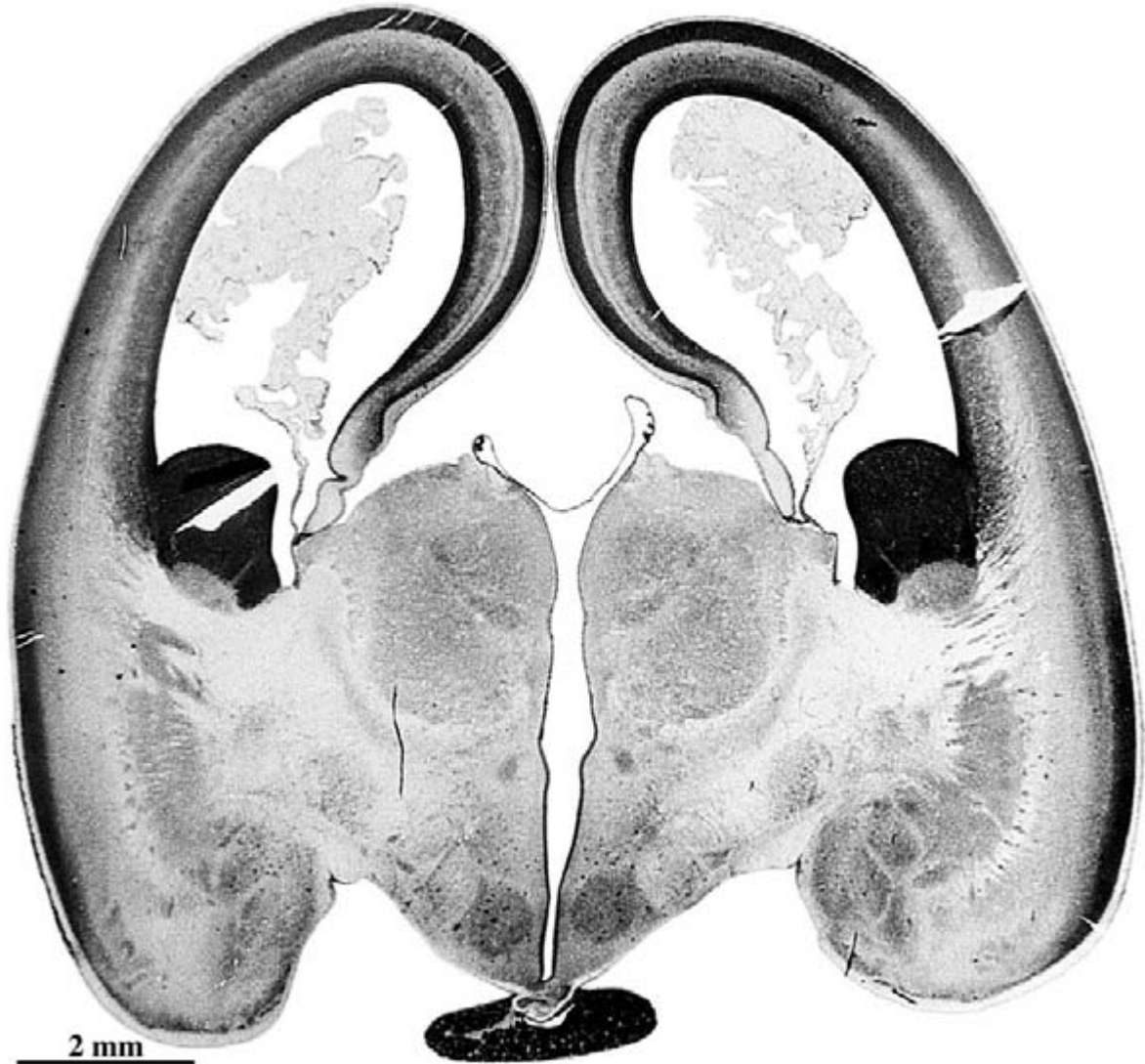
↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 9A

**GW11 Coronal
CR 60 mm, Y1-59
Level 9: Section 419**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

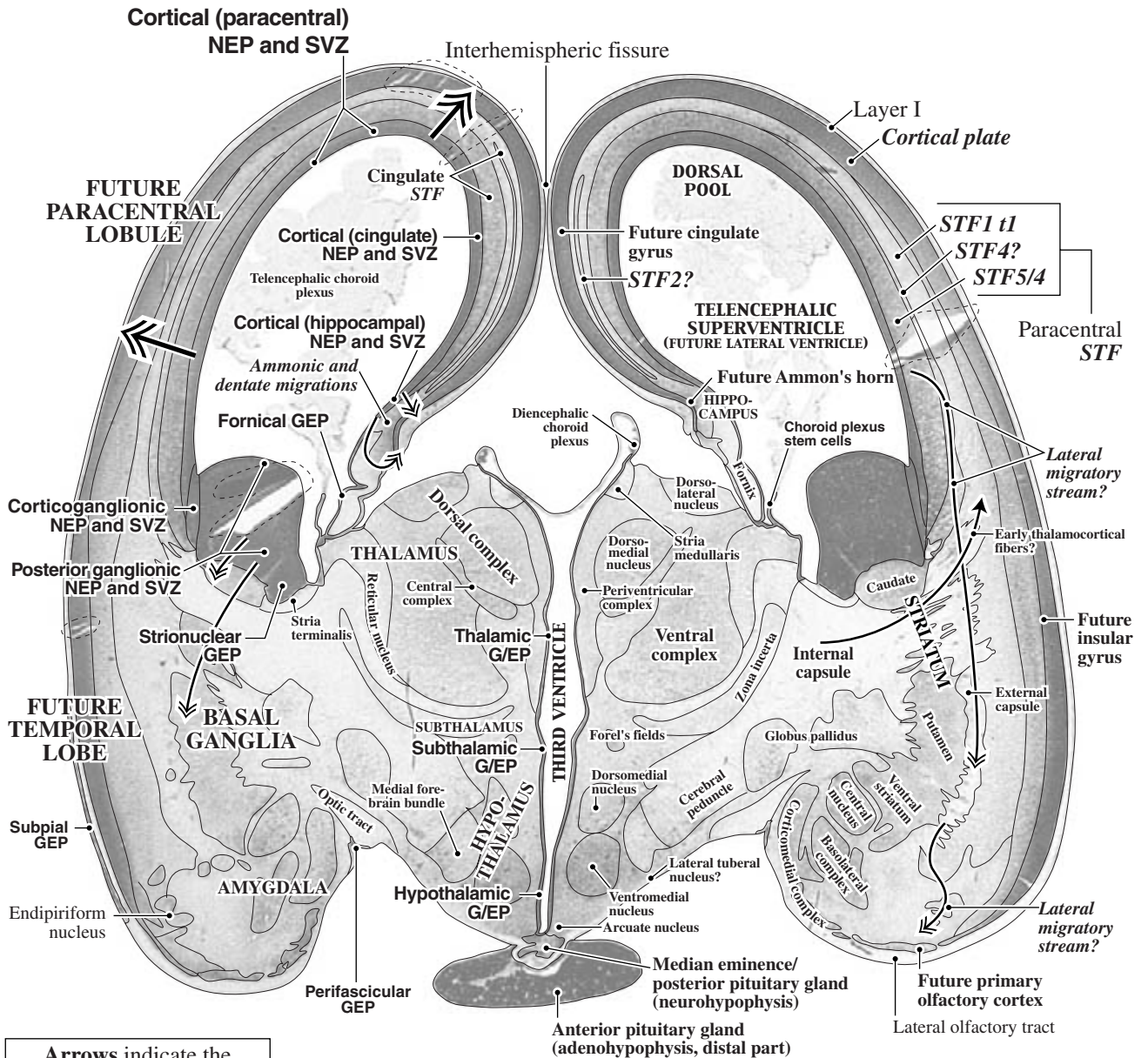
STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the diencephalon and basal telencephalon from level 9 in Plates 27A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

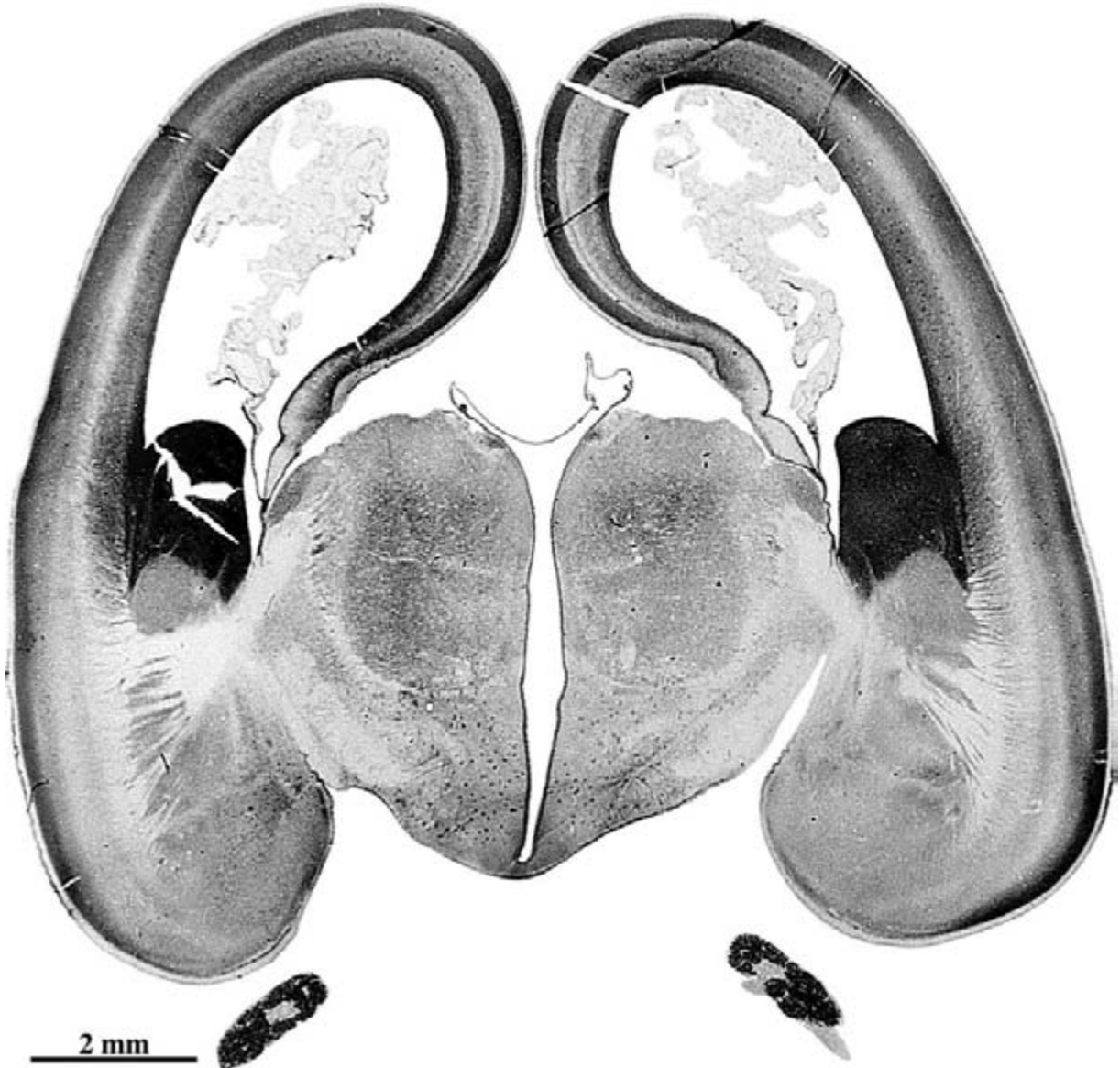
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 10A

**GW11 Coronal
CR 60 mm, Y1-59
Level 10: Section 439**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

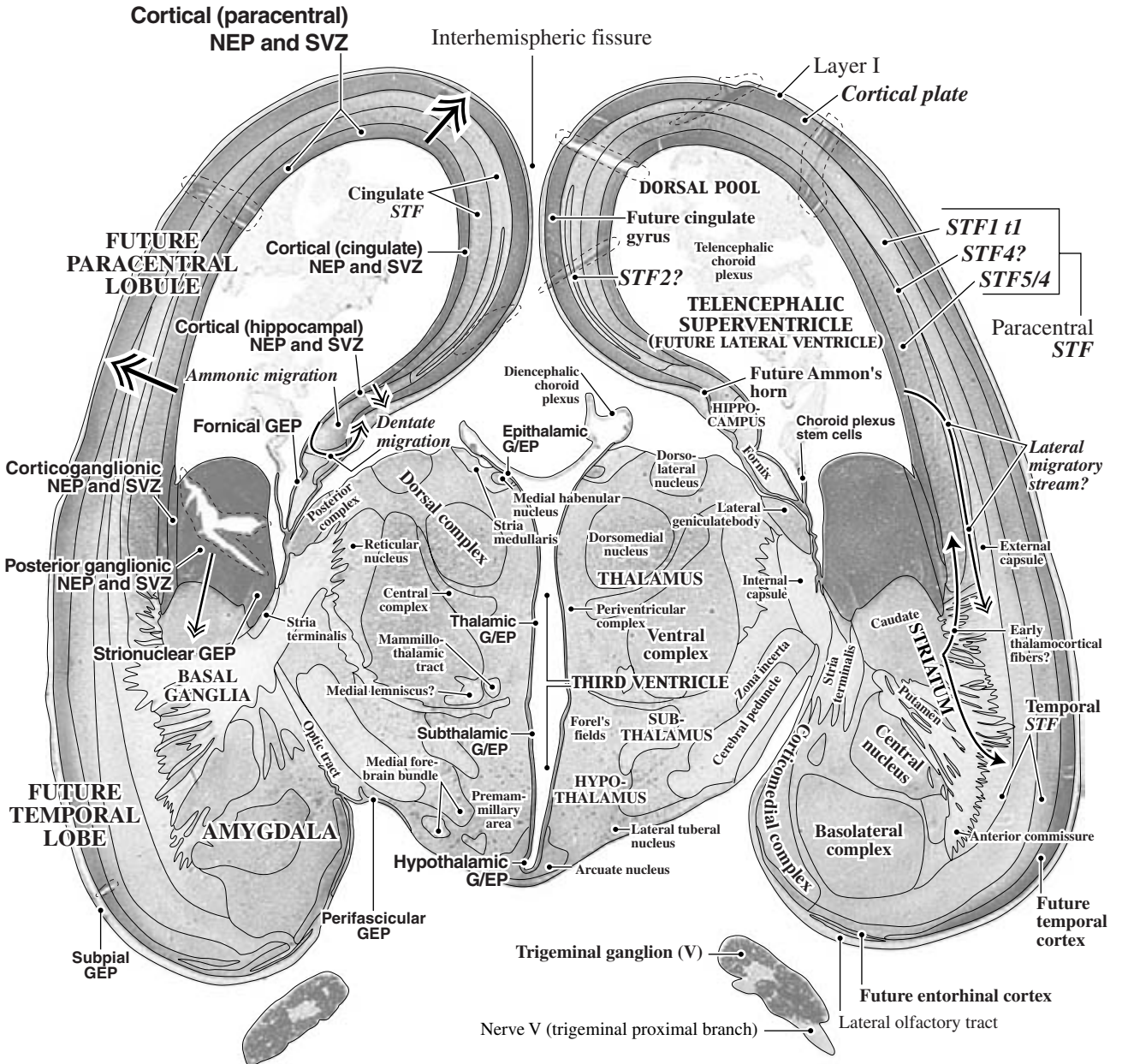
STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the diencephalon from section 449 in Plates 28A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
GEP - Helvetic bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Helvetic bold
G/EP - Helvetic bold
NEP - Helvetic bold
SVZ - Helvetic bold



↑ **Arrows indicate the presumed direction of axon growth** in brain fiber tracts.

↑ **Arrows indicate the presumed direction of neuron migration** from neuroepithelial sources.

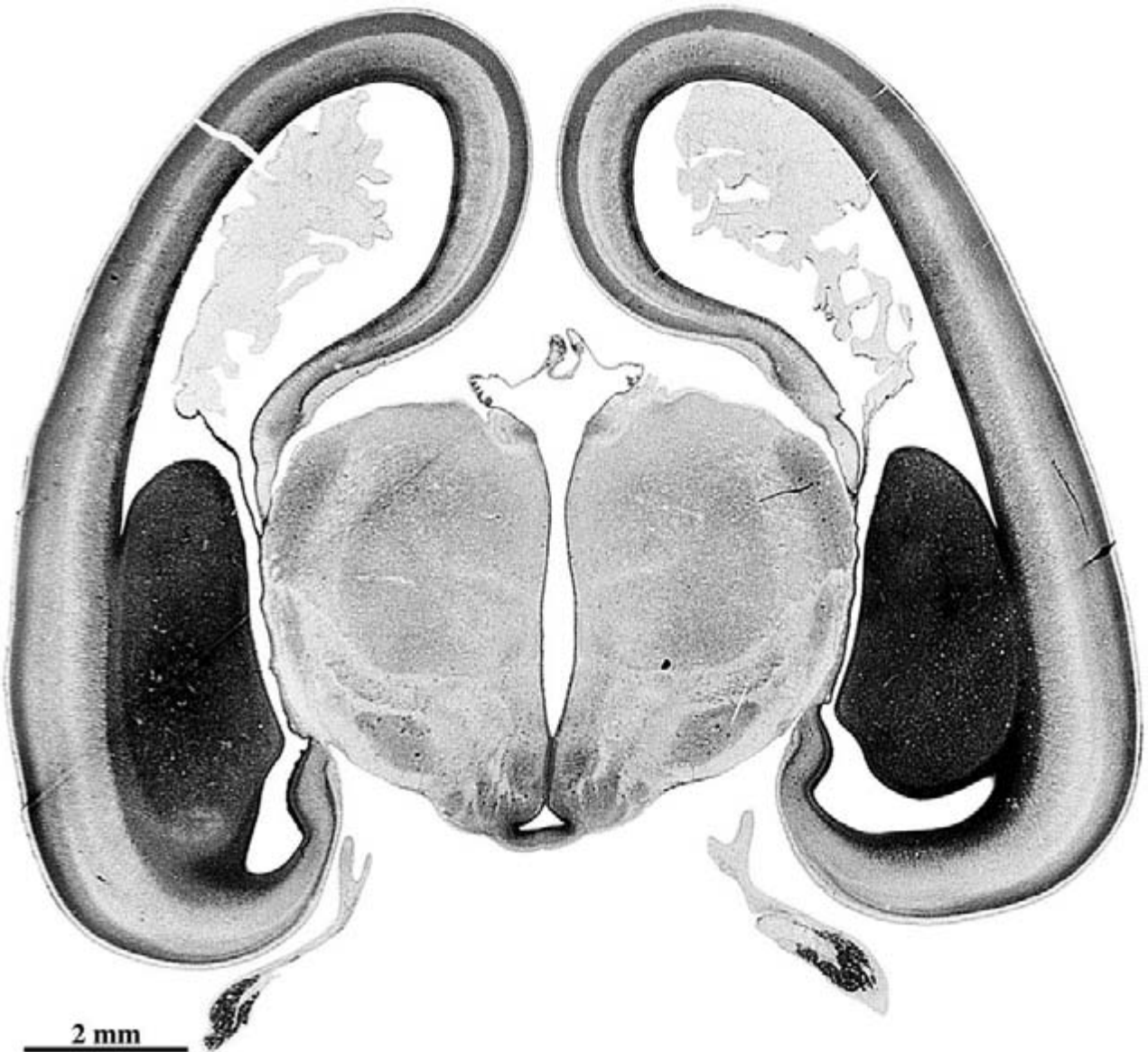
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 11A

**GW11 Coronal
CR 60 mm, Y1-59
Level 11: Section 459**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

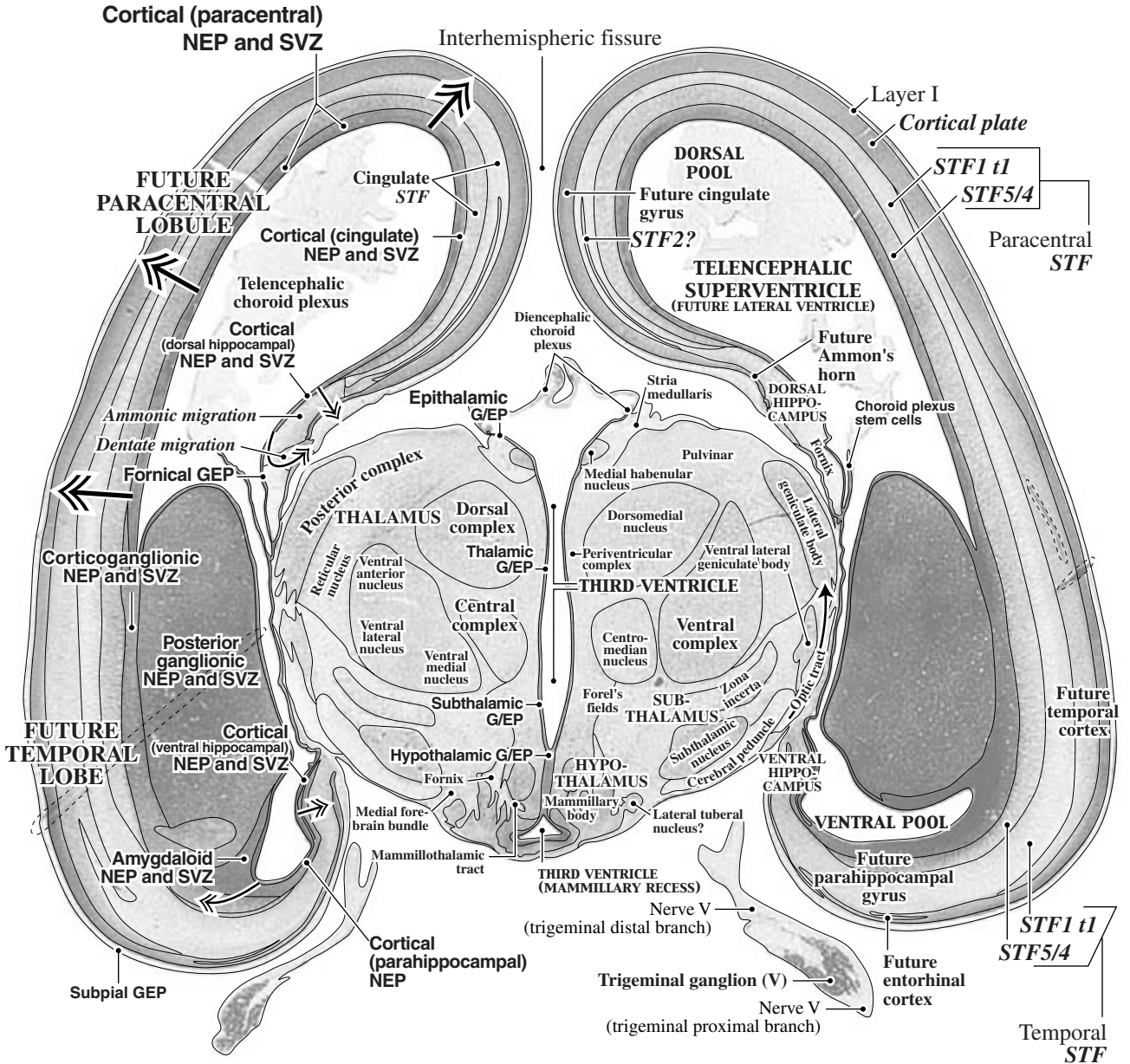
STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the diencephalon from section 449 in Plates 28A and B, from section 469 in Plates 29A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ **Arrows indicate the presumed direction of axon growth in brain fiber tracts.**

↑ **Arrows indicate the presumed direction of neuroepithelial migration from neuroepithelial sources.**

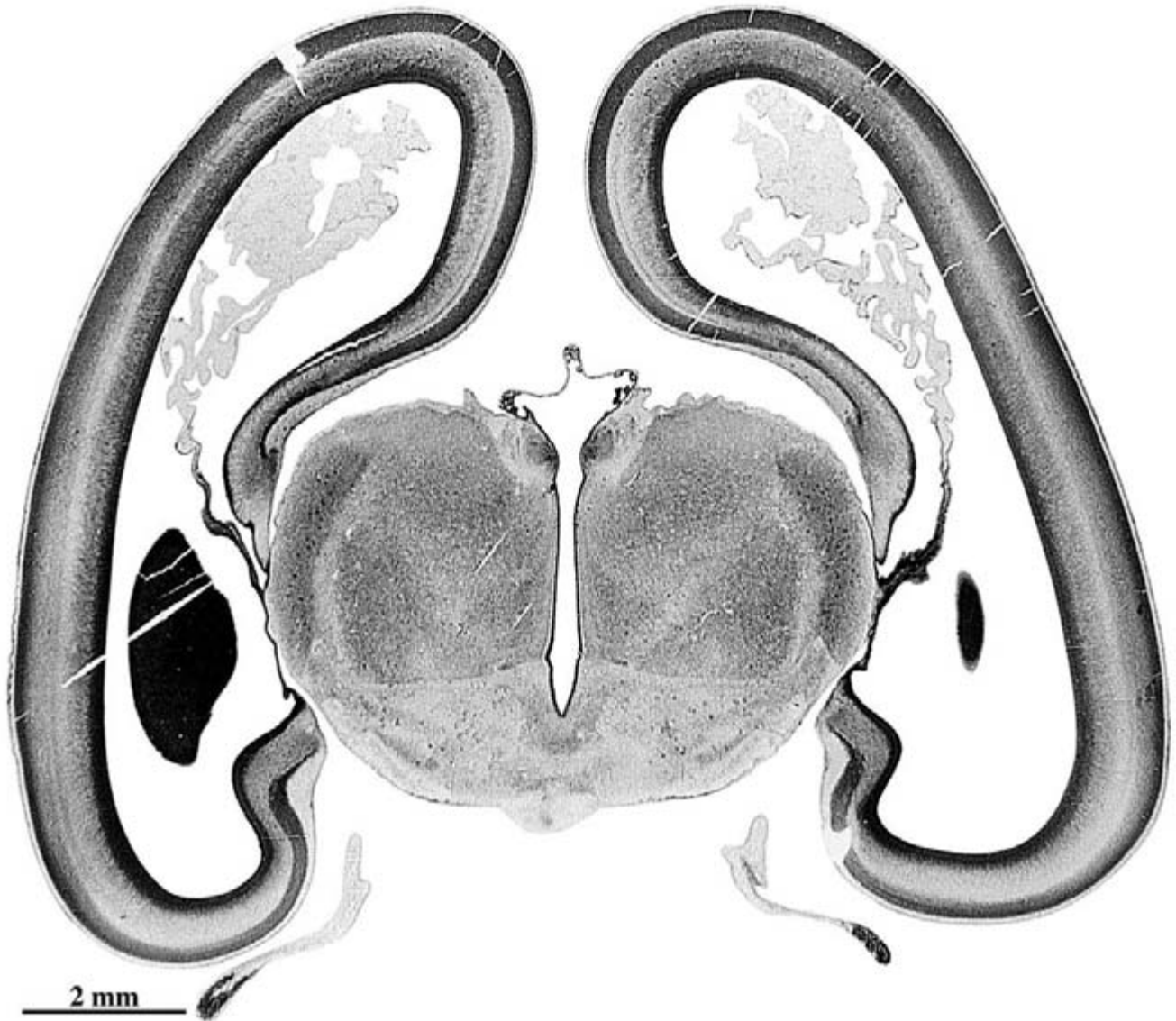
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 12A

**GW11 Coronal
CR 60 mm, Y1-59
Level 12: Section 479**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

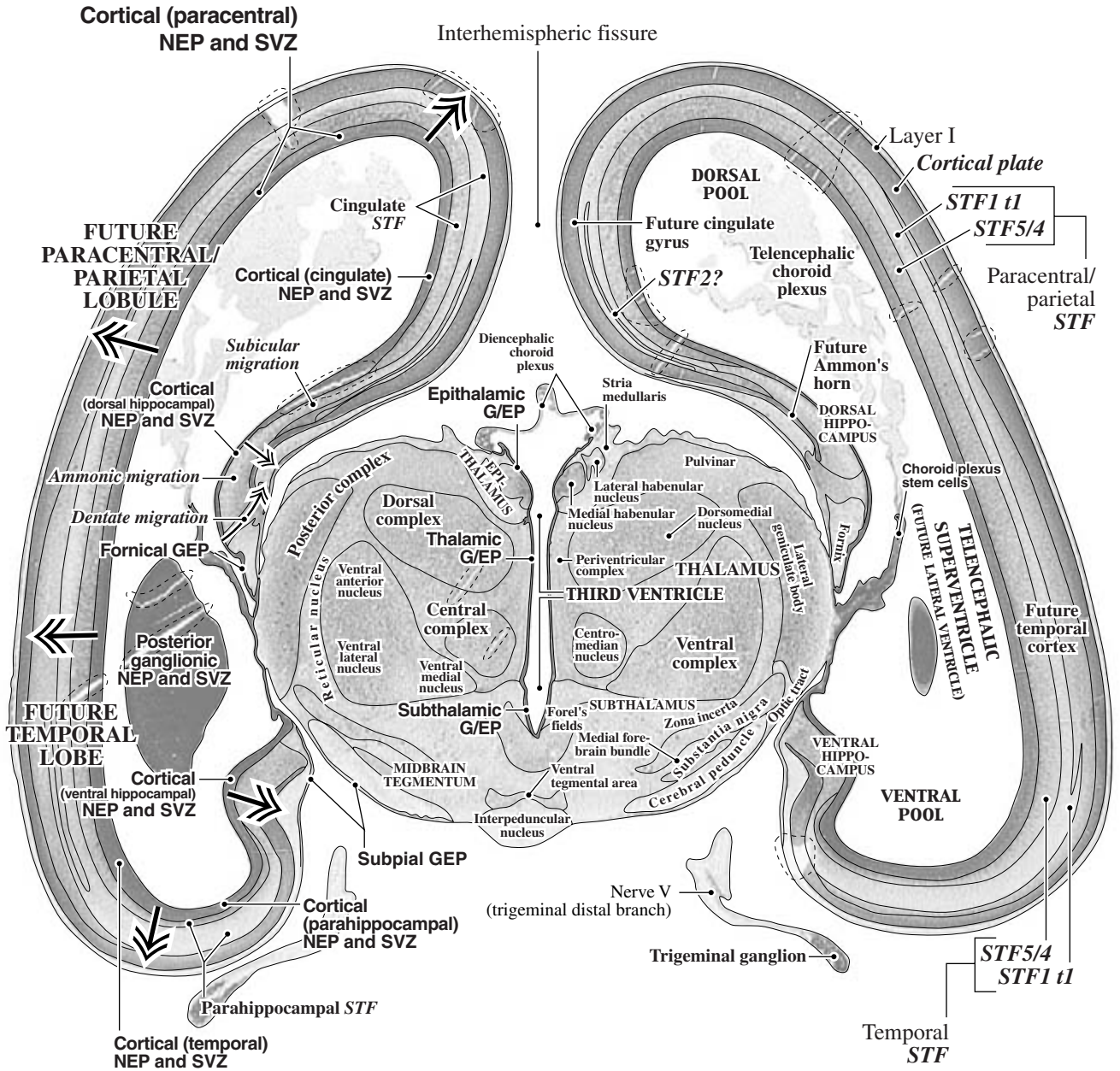
STF1 Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2 Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3 Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6 Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the diencephalon from section 469 in Plates 29A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

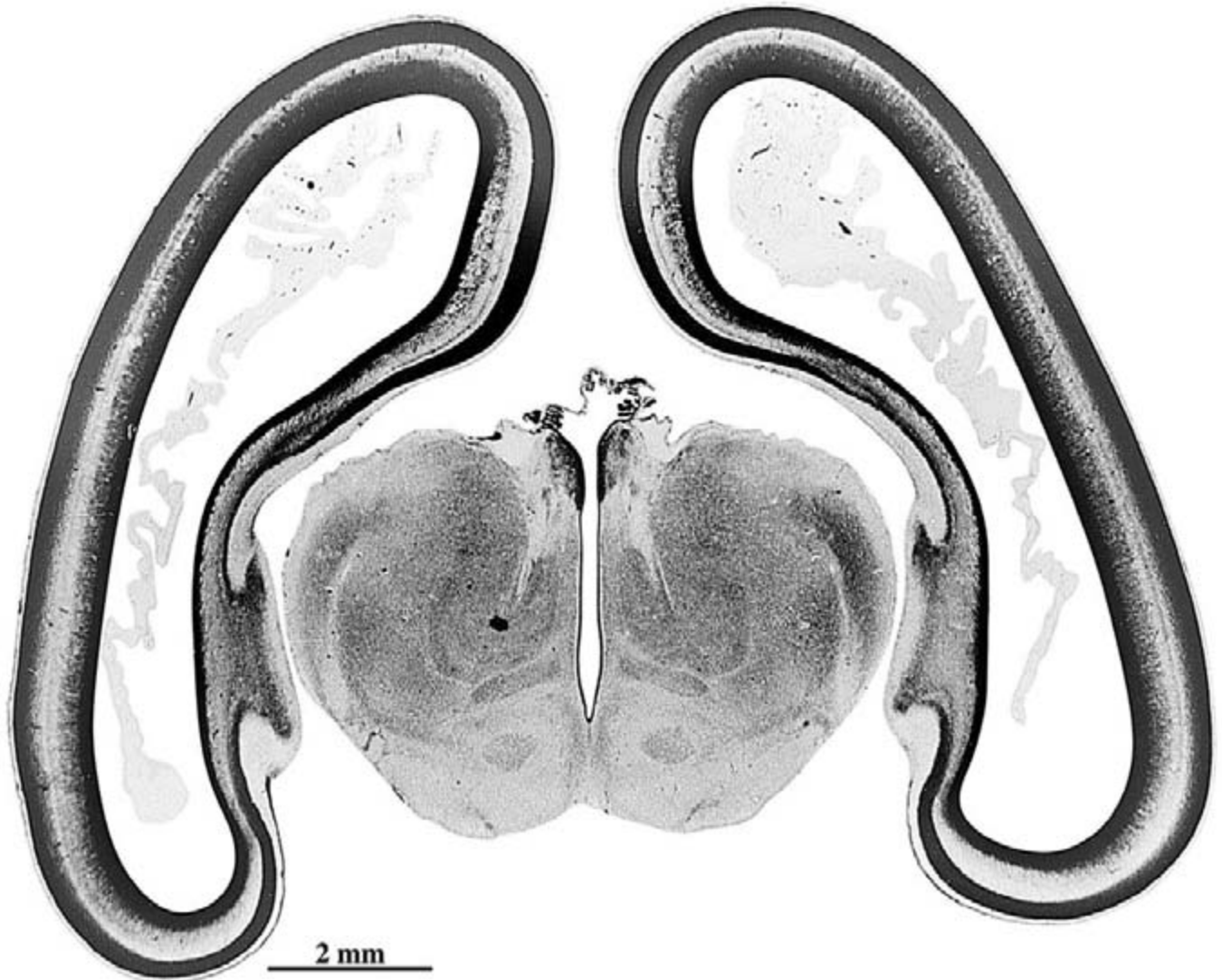
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 13A

**GW11 Coronal
CR 60 mm, Y1-59
Level 13: Section 500**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

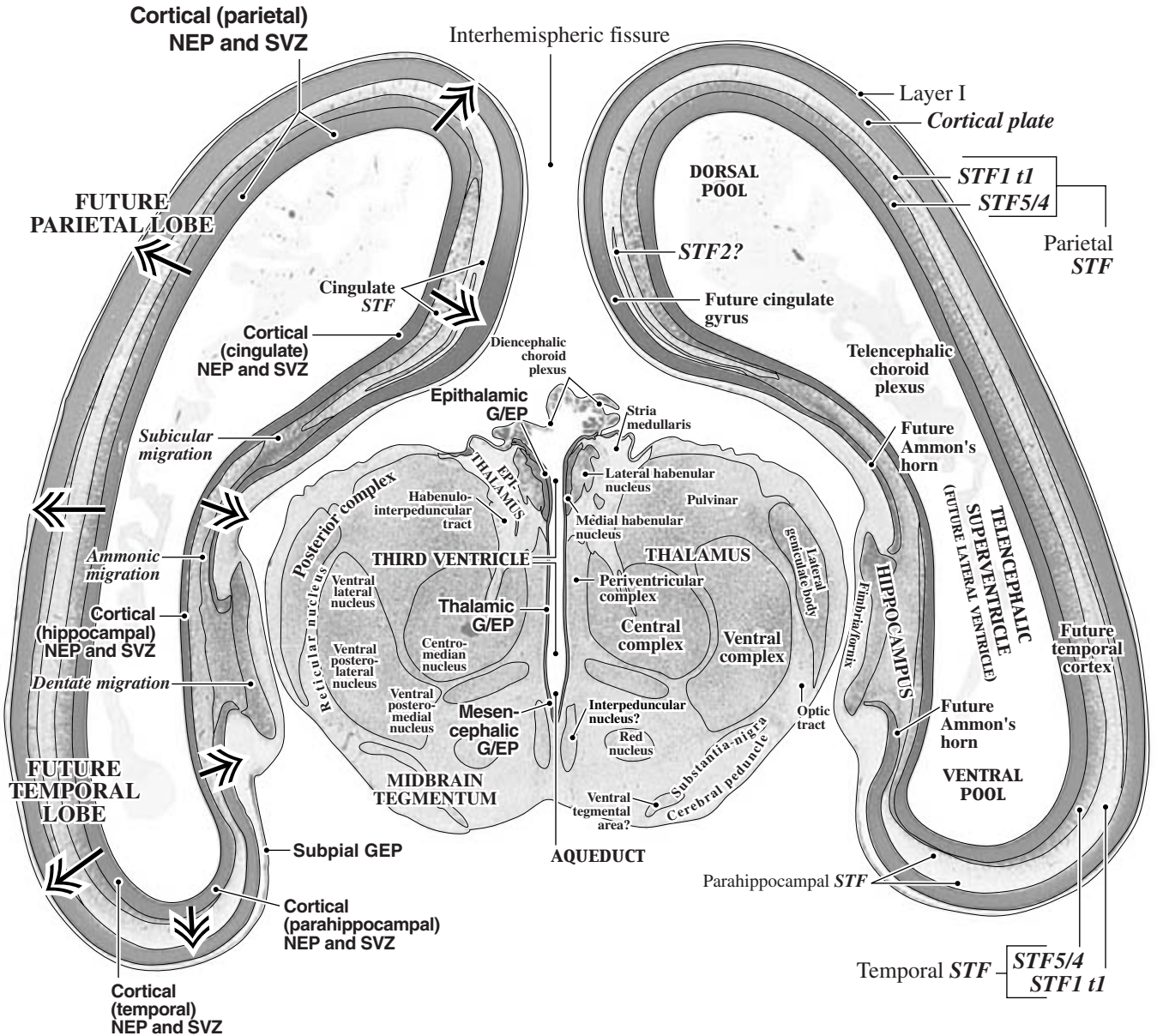
- | | | | |
|-------------|--|-------------|---|
| STF1 | Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter. | STF4 | Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons. |
| STF2 | Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate. | STF5 | Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix. |
| STF3 | Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices. | STF6 | Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester. |



See a high-magnification view of the midbrain and thalamus from section 499 in Plates 30A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

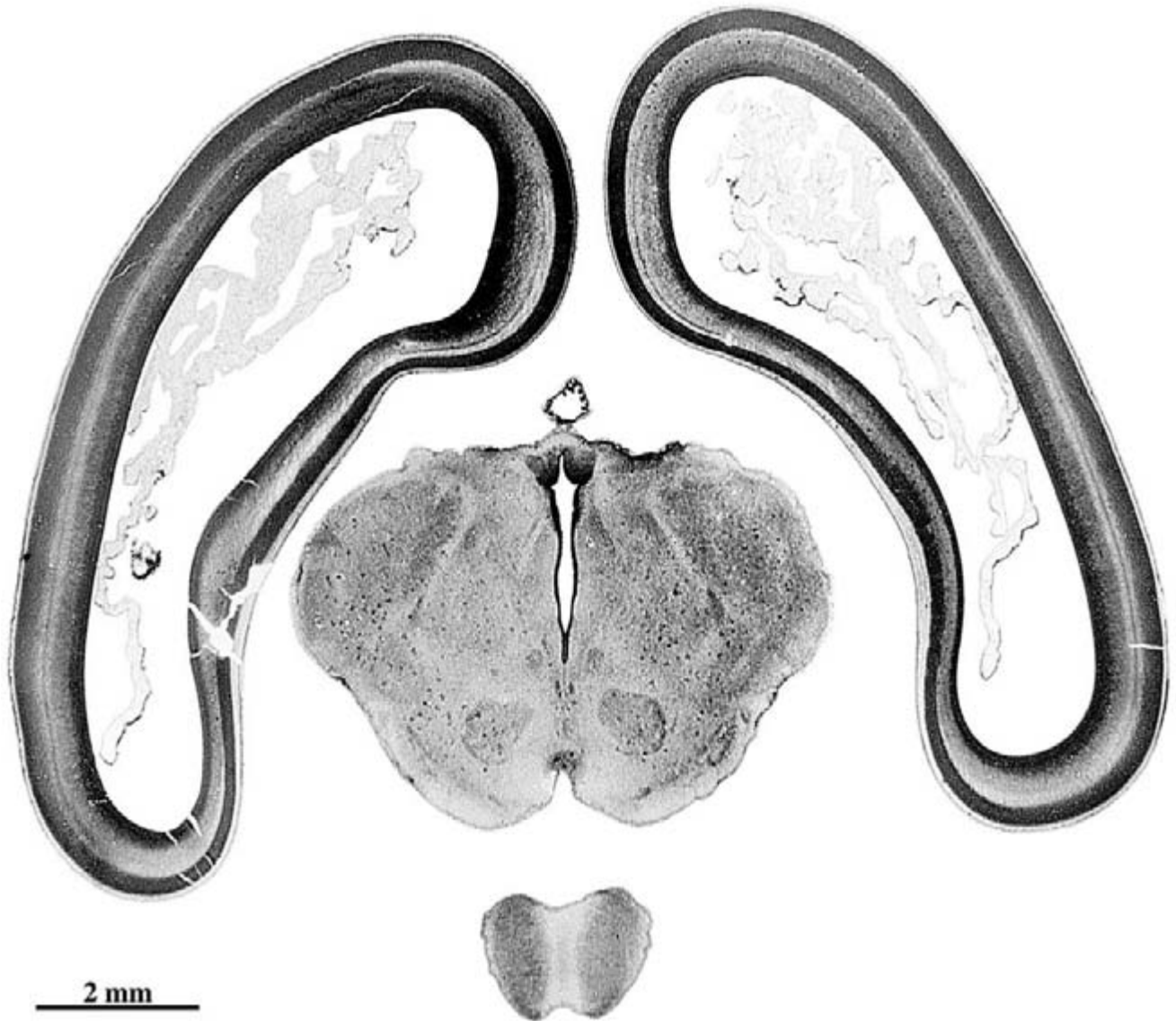
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 14A

**GW11 Coronal
CR 60 mm, Y1-59
Level 14: Section 529**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

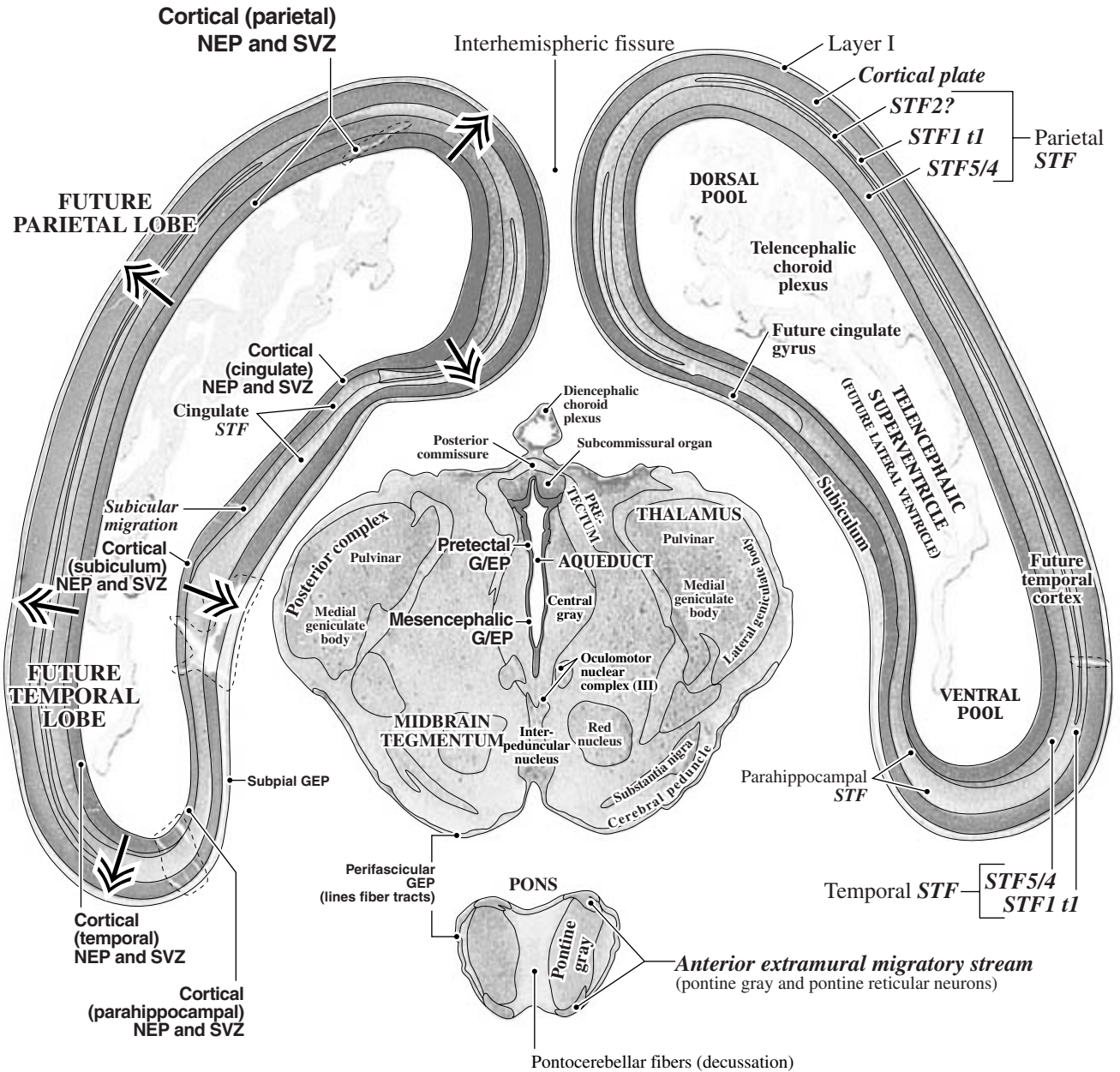
STF1	Superficial fibrous layer with an early developmental stage (<i>11</i>) when many cells are migrating through it, followed by a late stage (<i>12</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the parietal cortex from section 519 in Plates 23A and B, and of the thalamus and midbrain from this section in Plates 31A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - *Helvetica bold*
 Transient structure - *Times bold italic*
 Permanent structure - *Times Roman or Bold*

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

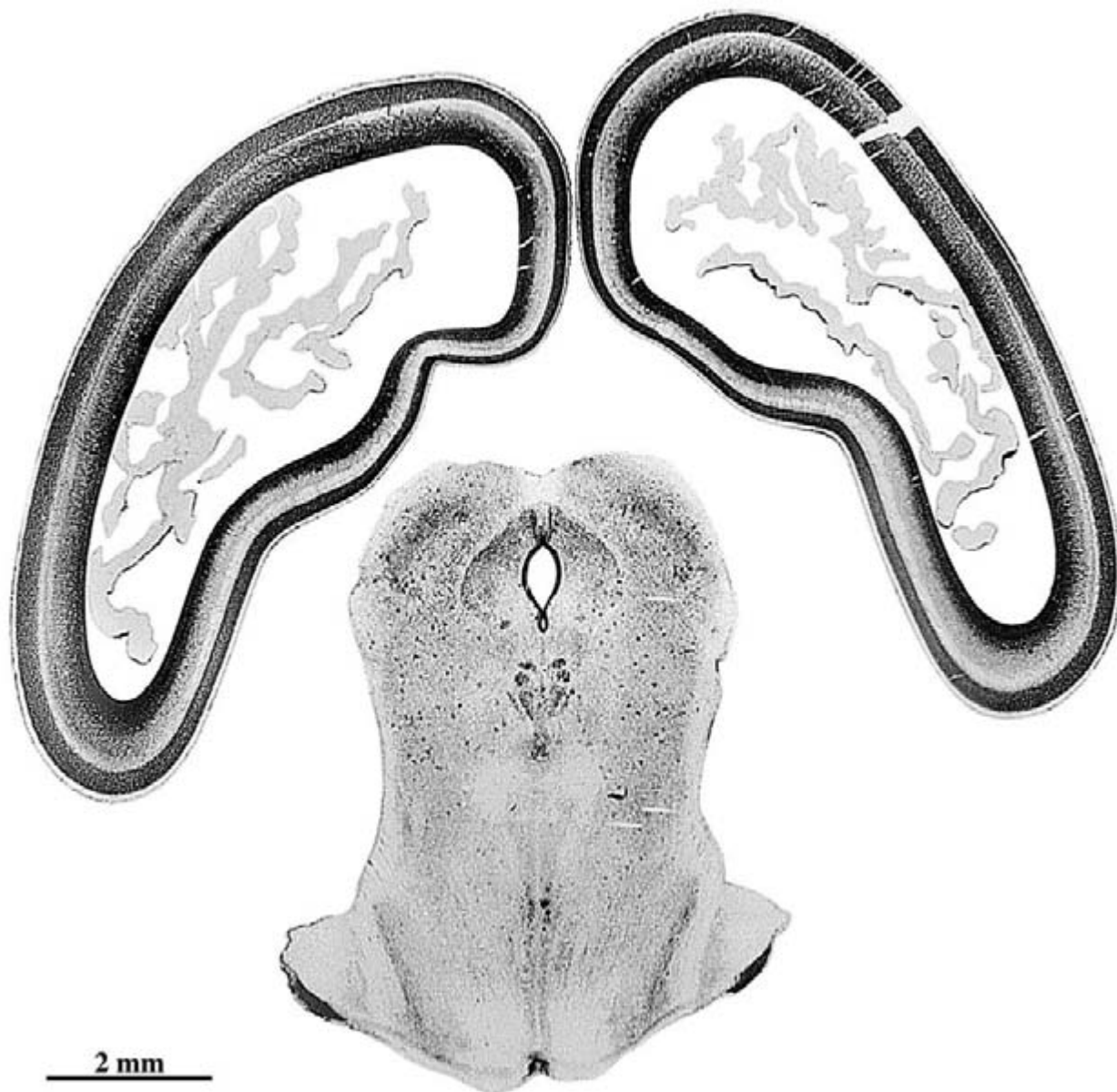
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 15A

**GW11 Coronal
CR 60 mm, Y1-59
Level 15: Section 569**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

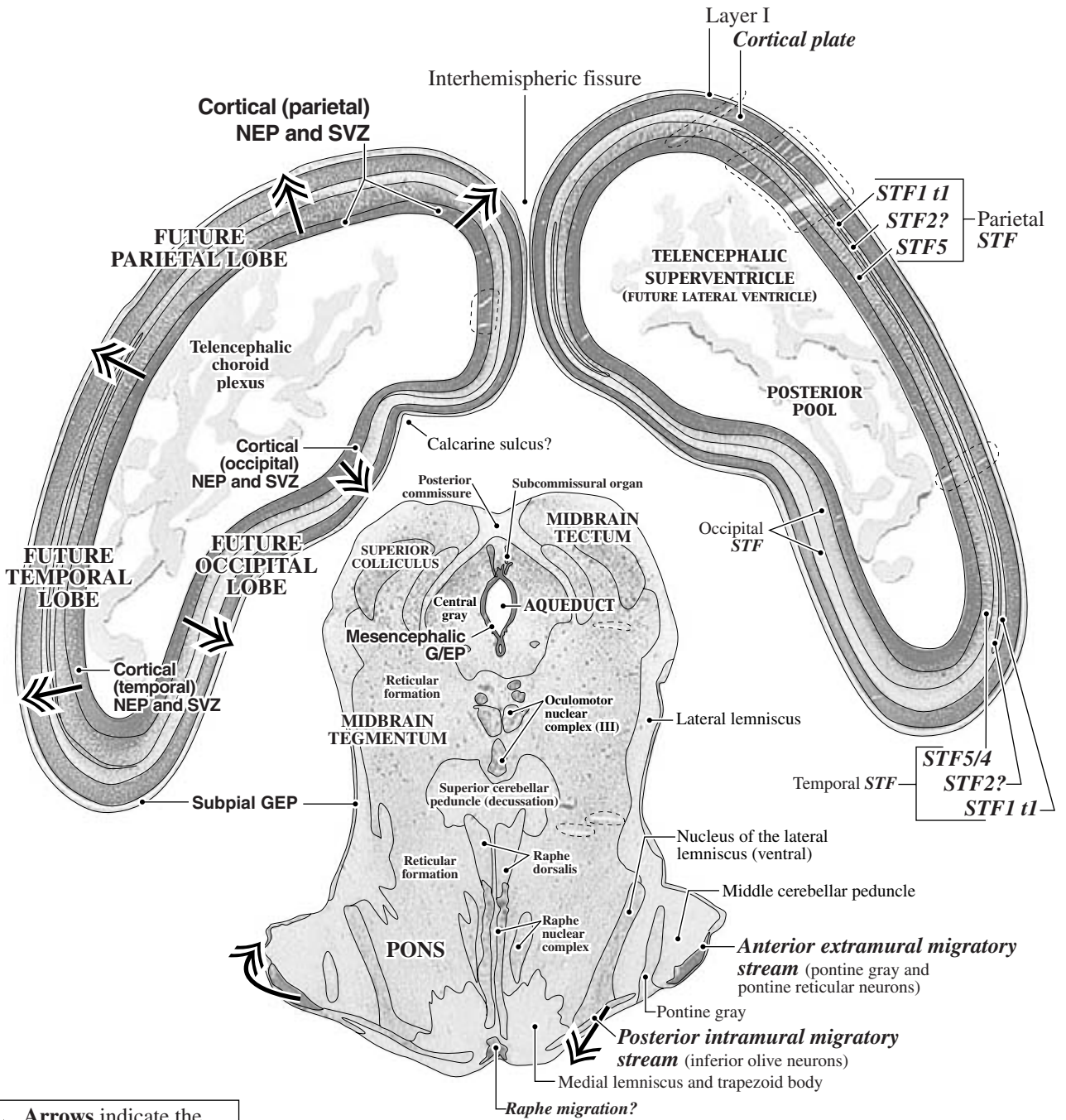
STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the midbrain and pons from section 549 in Plates 32A and B, and from this section in Plates 33A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 16A

**GW11 Coronal
CR 60 mm, Y1-59
Level 16: Section 599**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

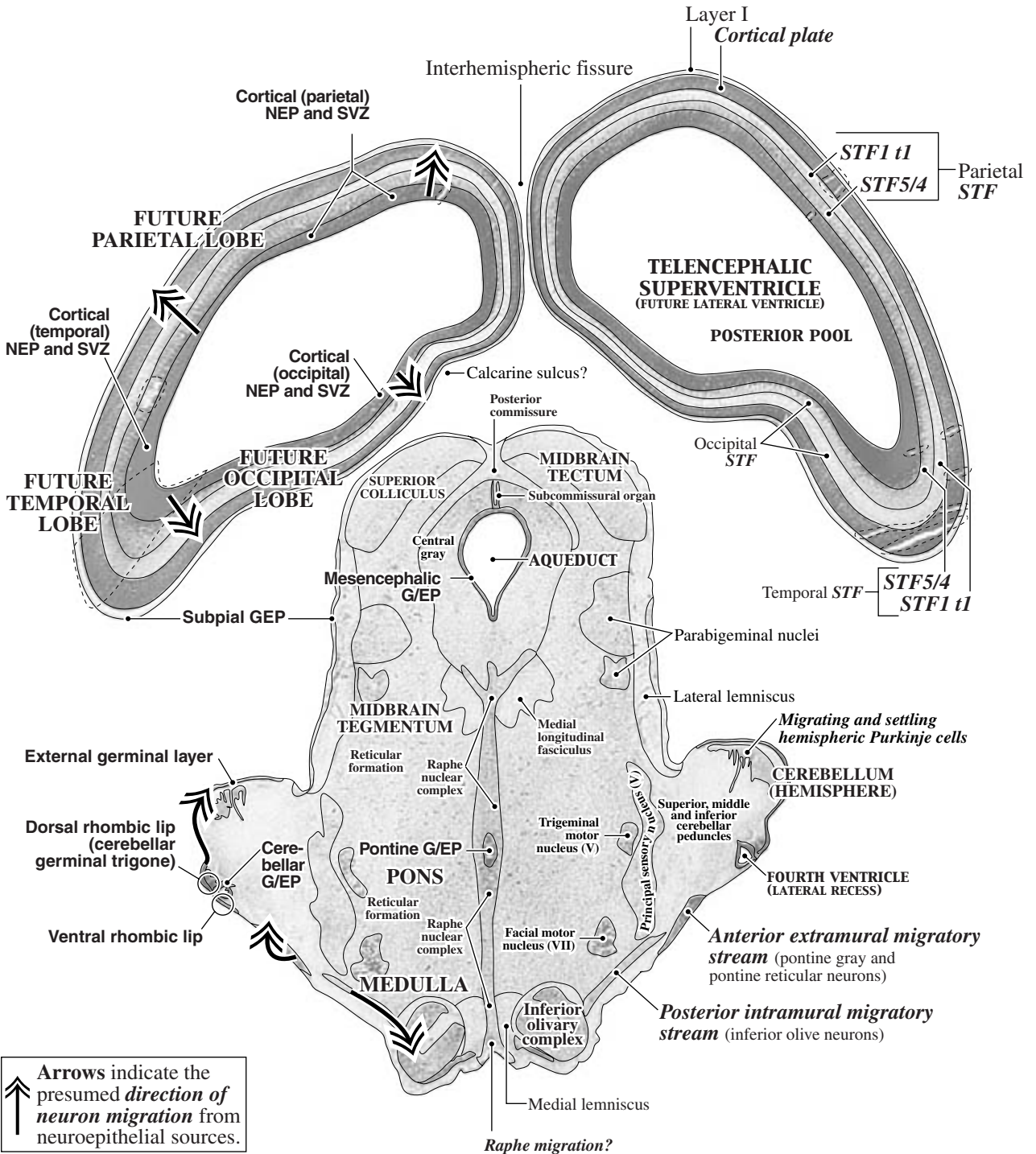
STF1	Superficial fibrous layer with an early developmental stage (<i>11</i>) when many cells are migrating through it, followed by a late stage (<i>12</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



See a high-magnification view of the midbrain and pons from section 589 in Plates 34A and B and from this section in Plates 35A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

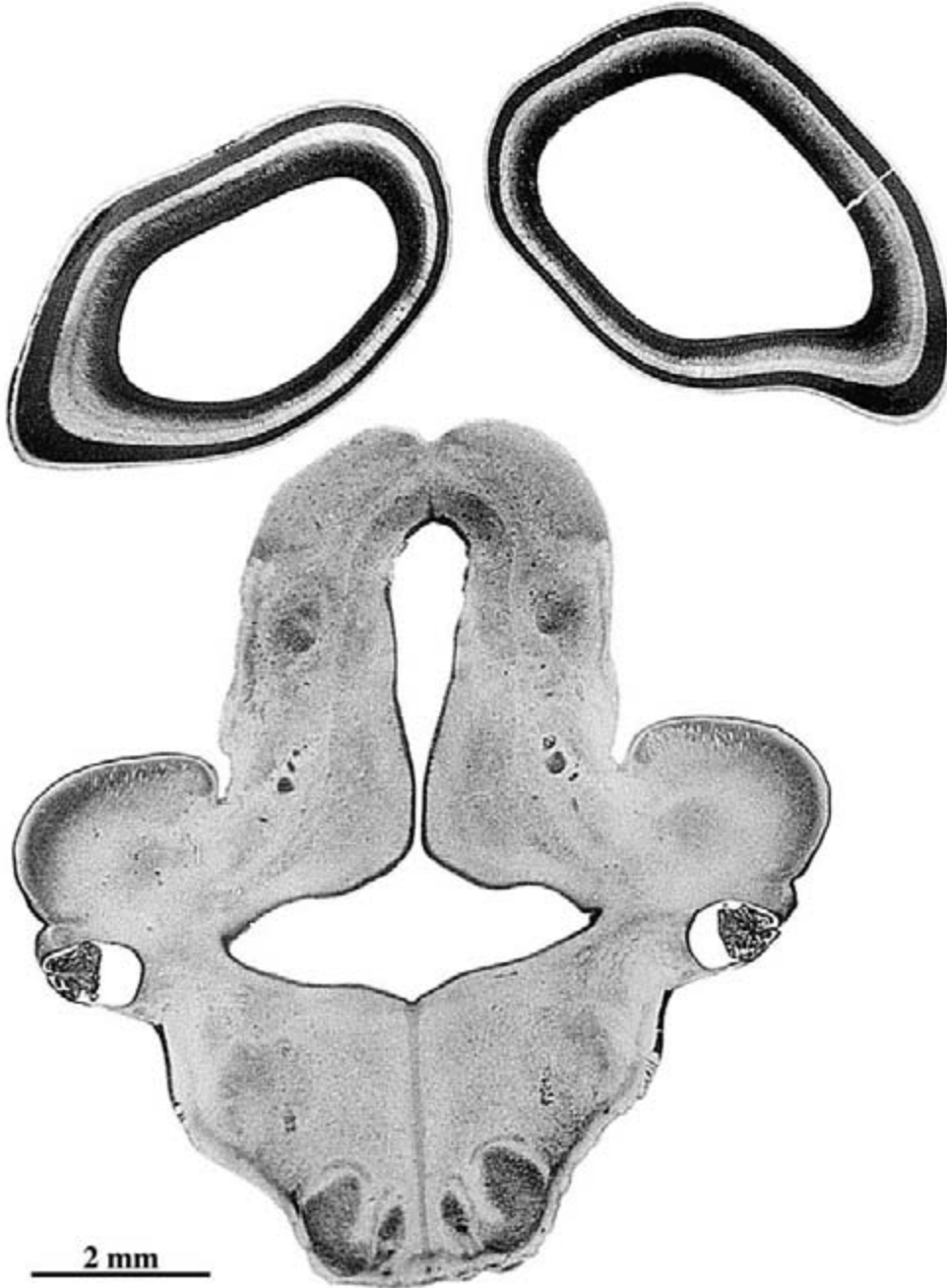
Dashed lines indicate staining and/or sectioning artifacts.

PLATE 17A

**GW11 Coronal
CR 60 mm, Y1-59
Level 17: Section 629**

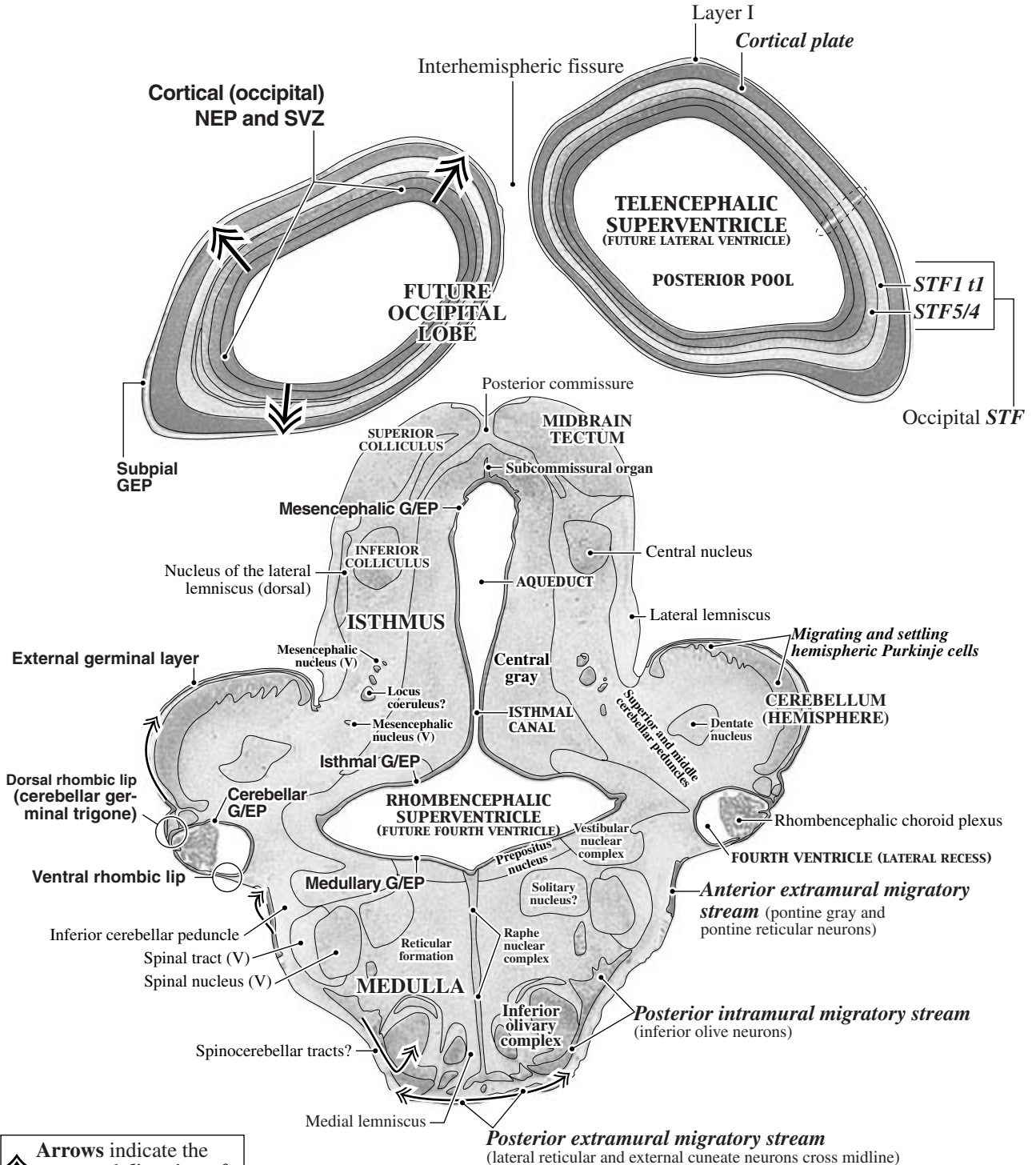
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 18A

**GW11 Coronal
CR 60 mm, Y1-59
Level 18: Section 649**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>t1</i>) when many cells are migrating through it, followed by a late stage (<i>t2</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

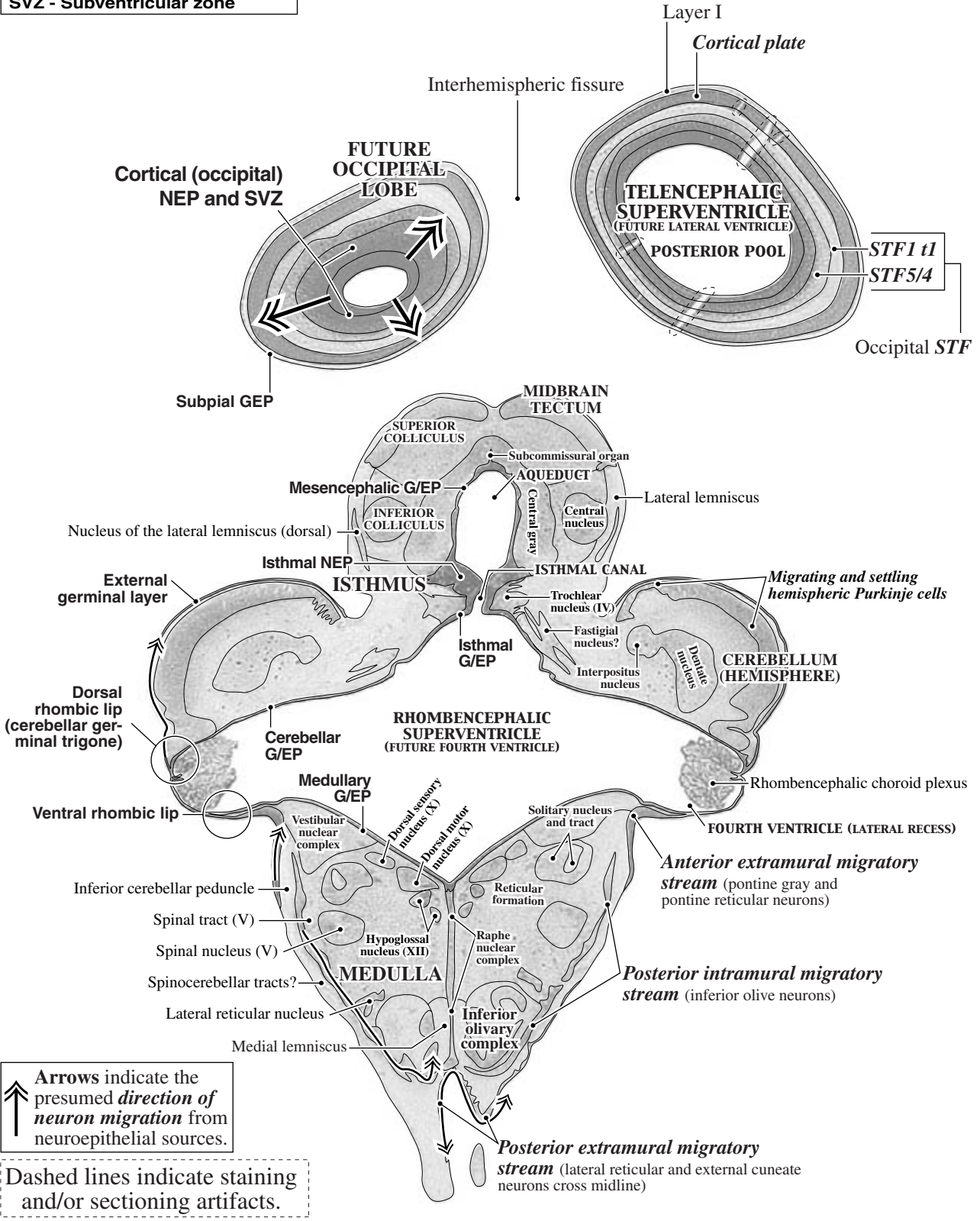


PLATE 19A

**GW11 Coronal
CR 60 mm, Y1-59
Level 19: Section 659**

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1	Superficial fibrous layer with an early developmental stage (<i>11</i>) when many cells are migrating through it, followed by a late stage (<i>12</i>) with sparse cells. Endures as the subcortical white matter.	STF4	Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
STF2	Upper cellular layer, the most superficial sojourn zone where cells translocate to the cortical plate.	STF5	Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.
STF3	Honeycomb trilaminar matrix (<i>3a, 3b, 3c</i>) of cells and fibers that is not present during the first trimester and is found in granular (sensory) cortices.	STF6	Late-forming deep layer of callosal fibers outside the germinal matrix that is not present during the first trimester.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
GEP - Glioeptihelium
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioeptihelium
G/EP - Glioeptihelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

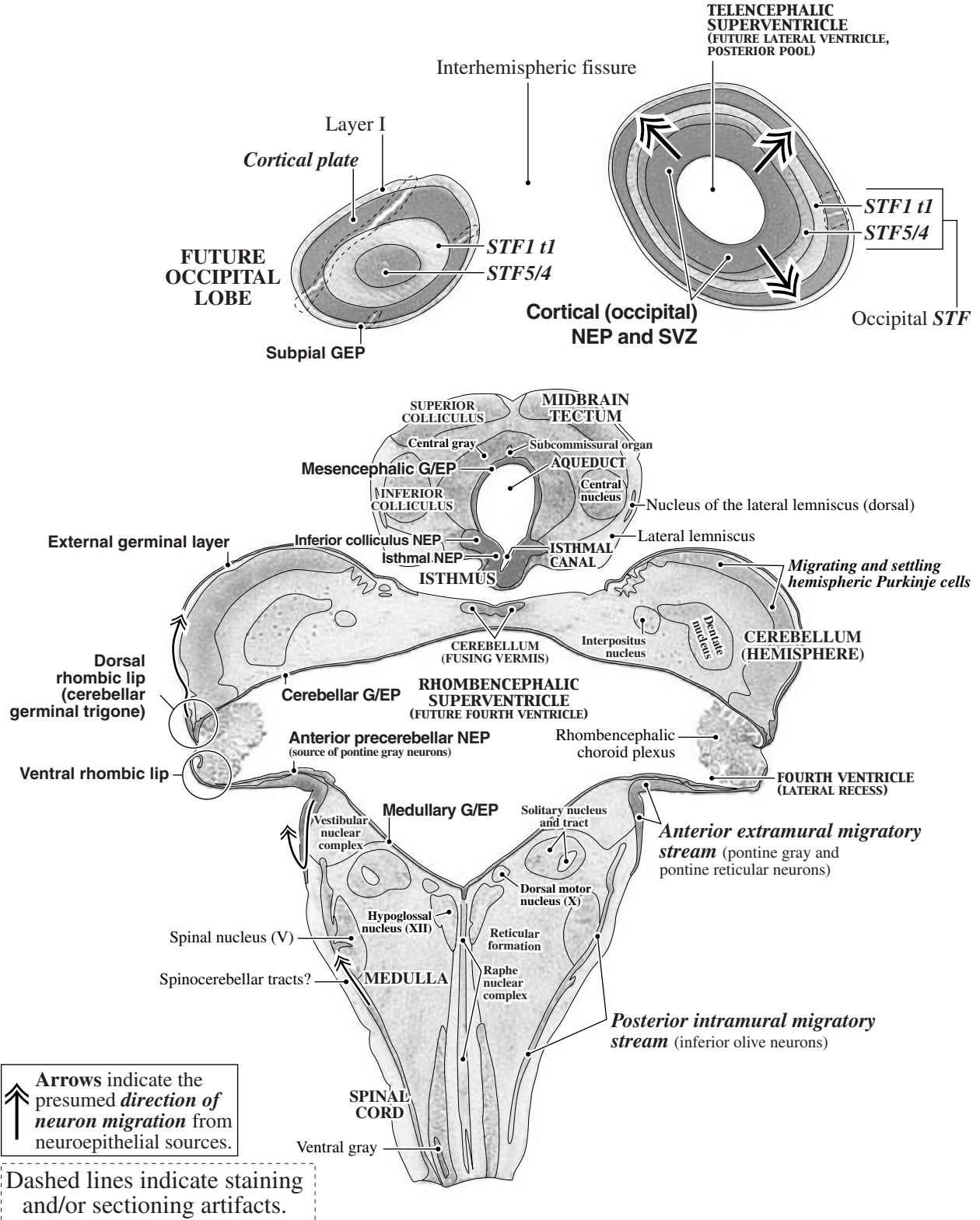
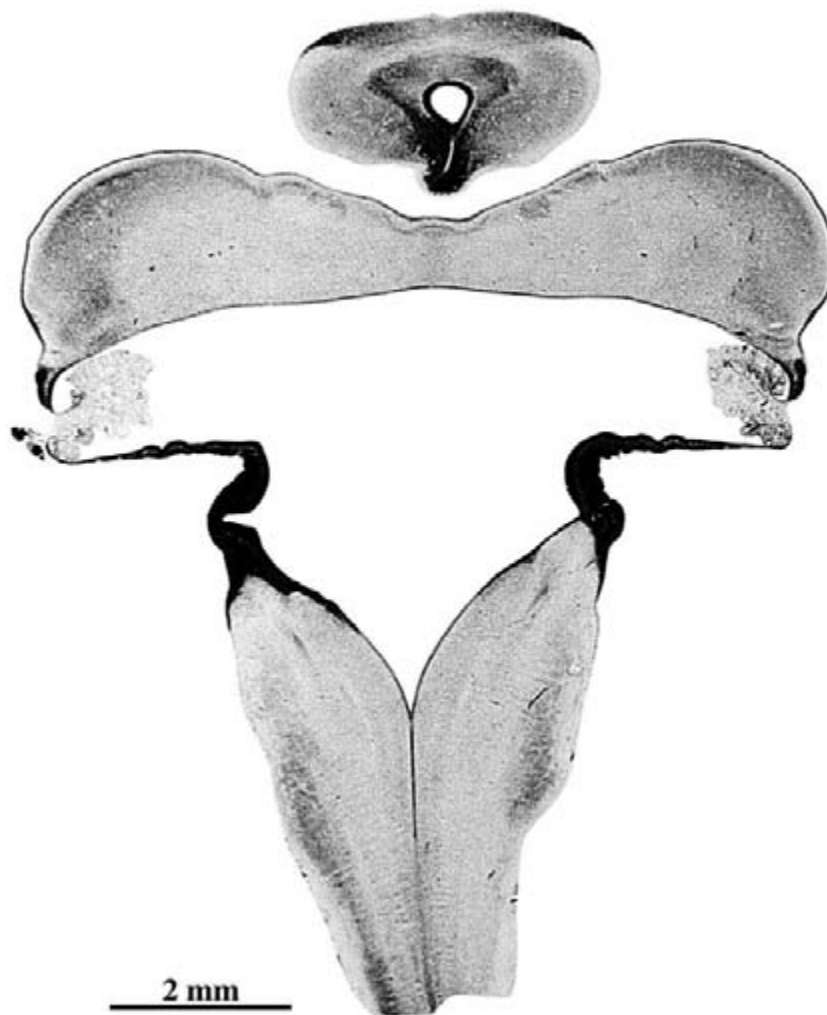


PLATE 20A

**GW11 Coronal
CR 60 mm, Y1-59
Level 20: Section 680**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

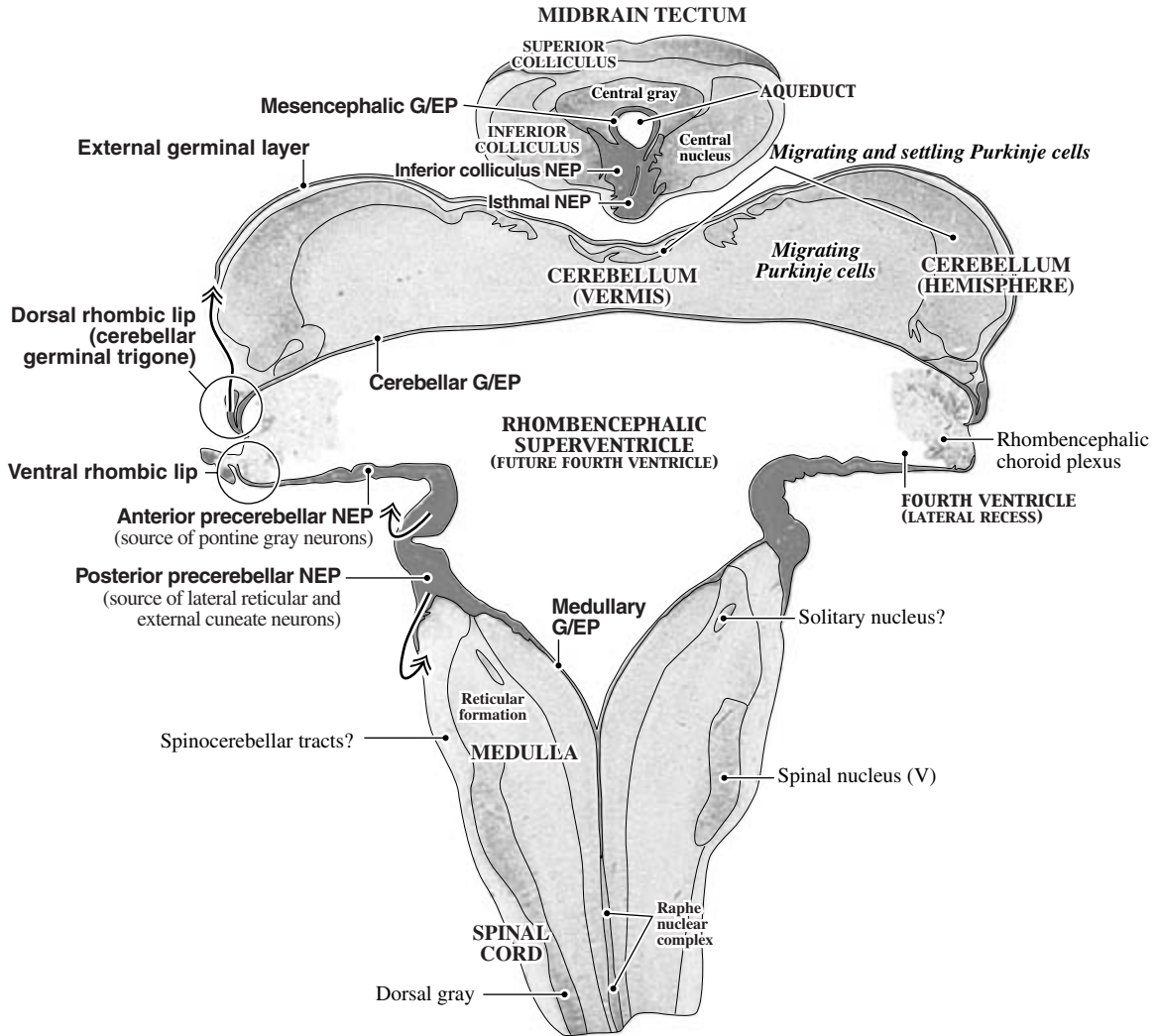
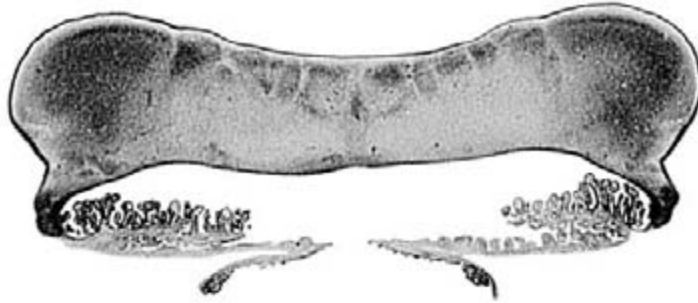


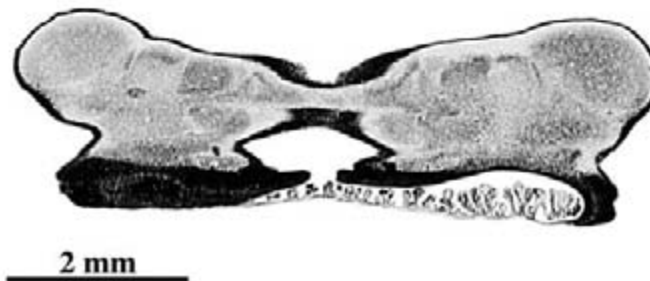
PLATE 21A

**GW11 Coronal
CR 60 mm, Y1-59**

Level 21: Section 709



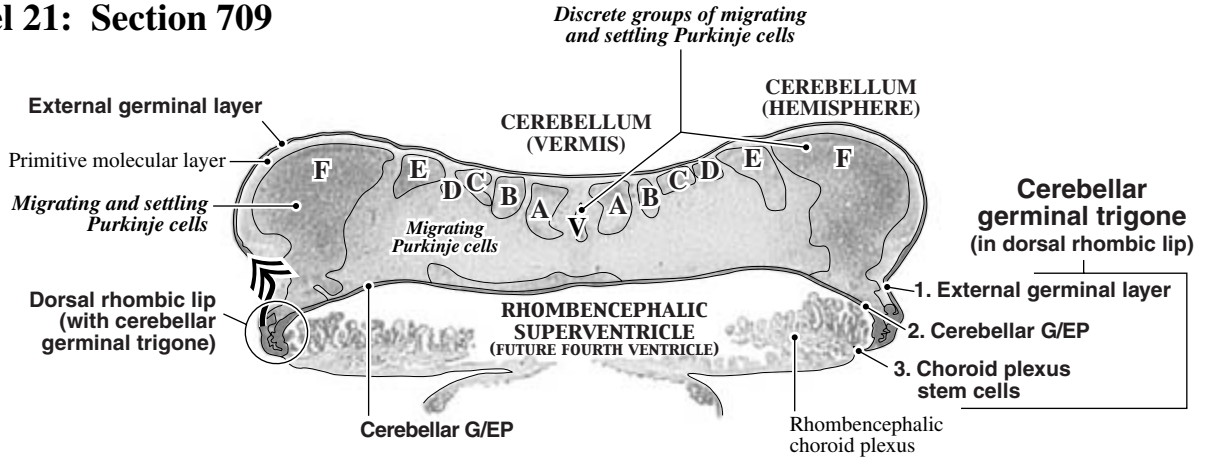
Level 22: Section 720



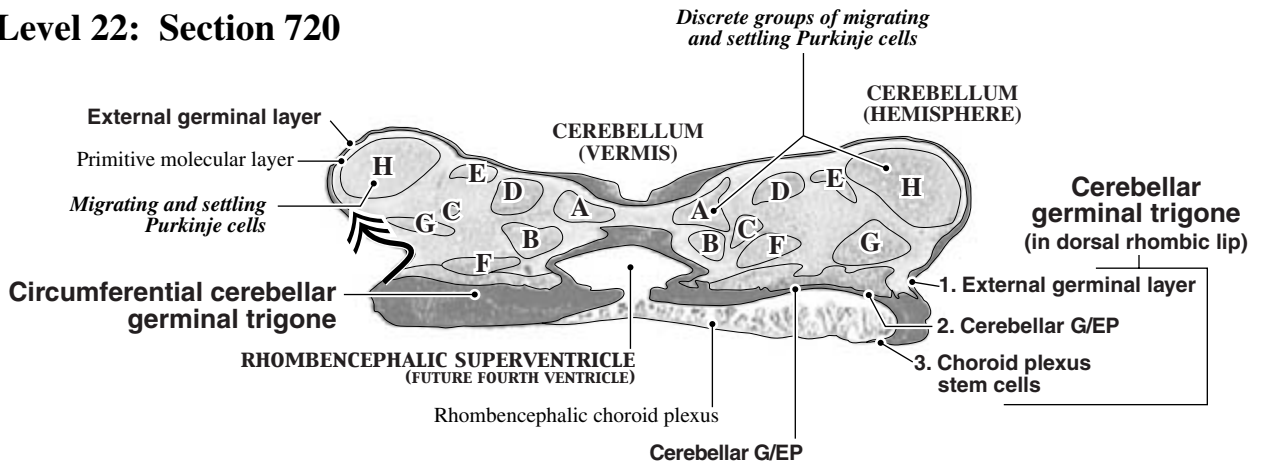
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

Gioepithelium/ependyma - G/EP

Level 21: Section 709



Level 22: Section 720



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 22A

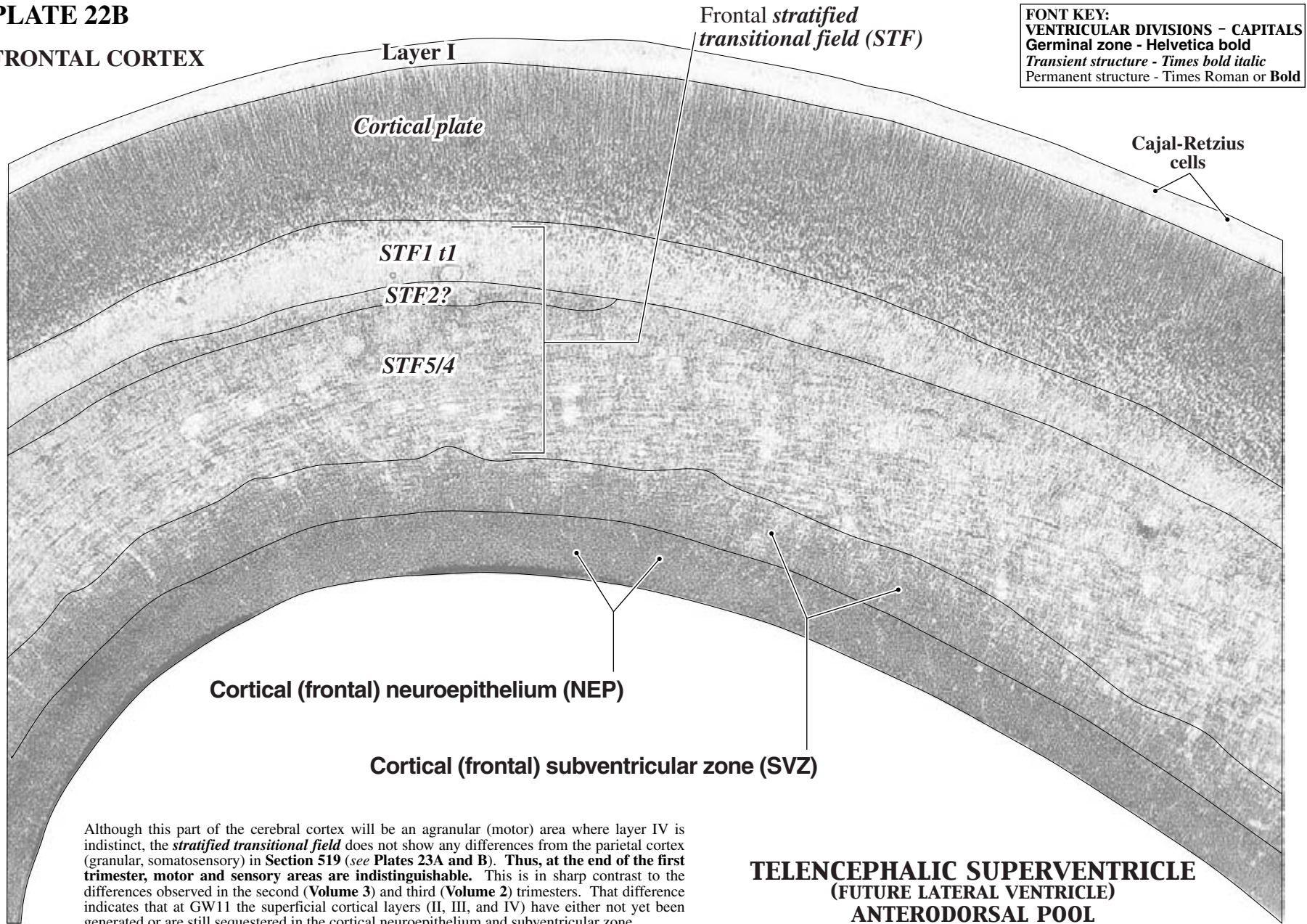
**GW11 Coronal
CR 60 mm, Y1-59
Between levels 2 and 3:
Section 269
FRONTAL
CORTEX**



See level 2 in Plates 2A and B, level 3 in Plates 3A and B.

PLATE 22B

FRONTAL CORTEX



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

Although this part of the cerebral cortex will be an agranular (motor) area where layer IV is indistinct, the *stratified transitional field* does not show any differences from the parietal cortex (granular, somatosensory) in Section 519 (see Plates 23A and B). Thus, at the end of the first trimester, motor and sensory areas are indistinguishable. This is in sharp contrast to the differences observed in the second (Volume 3) and third (Volume 2) trimesters. That difference indicates that at GW11 the superficial cortical layers (II, III, and IV) have either not yet been generated or are still sequestered in the cortical neuroepithelium and subventricular zone.

**TELENCEPHALIC SUPERVENTRICLE
 (FUTURE LATERAL VENTRICLE)
 ANTERODORSAL POOL**

PLATE 23A

**GW11 Coronal
CR 60 mm, Y1-59
Near level 14:
Section 519
PARIETAL
CORTEX**

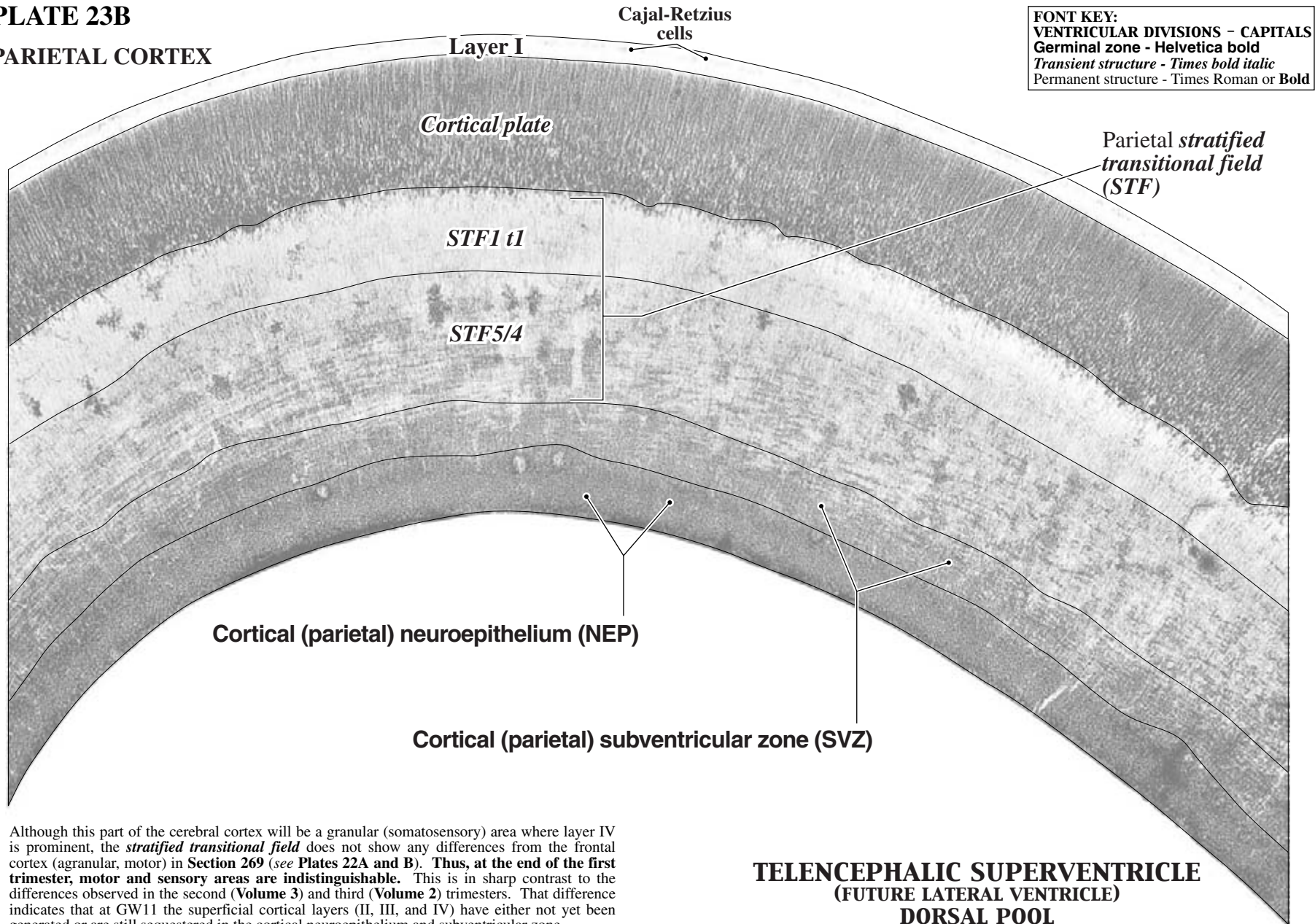


0.25 mm

See level 14 in Plates 14A and B.

PLATE 23B

PARIETAL CORTEX



Although this part of the cerebral cortex will be a granular (somatosensory) area where layer IV is prominent, the *stratified transitional field* does not show any differences from the frontal cortex (agranular, motor) in **Section 269** (see **Plates 22A and B**). **Thus, at the end of the first trimester, motor and sensory areas are indistinguishable.** This is in sharp contrast to the differences observed in the second (**Volume 3**) and third (**Volume 2**) trimesters. That difference indicates that at GW11 the superficial cortical layers (II, III, and IV) have either not yet been generated or are still sequestered in the cortical neuroepithelium and subventricular zone.

PLATE 24A

**GW11 Coronal
CR 60 mm, Y1-59
Between levels 8 and 9:
Section 399**

**PARACENTRAL
CORTEX**

**See levels 8 and 9
in Plates
8 to 9A and B.**

**Enlarged in
Plates 25A and B.**

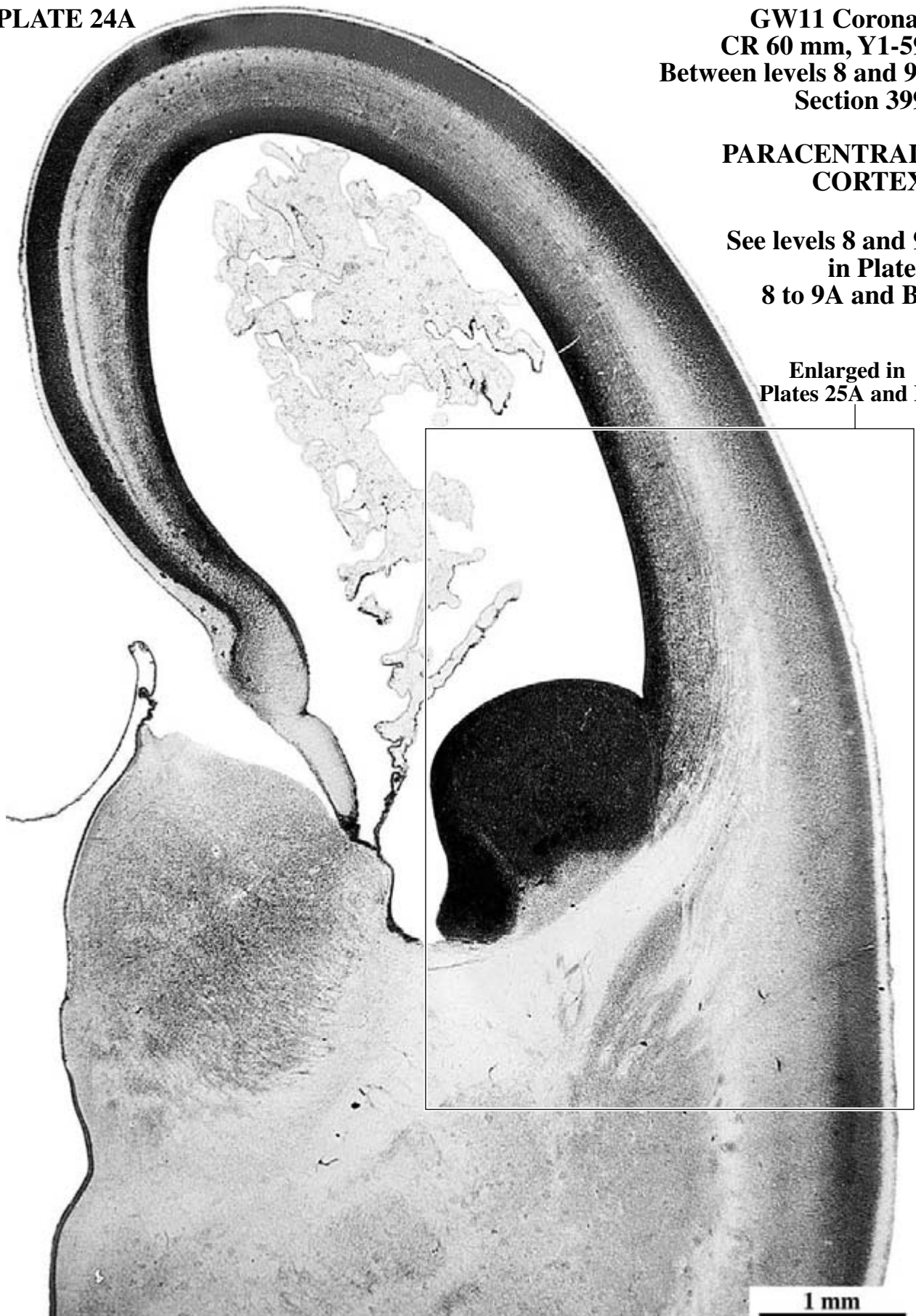


PLATE 24B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

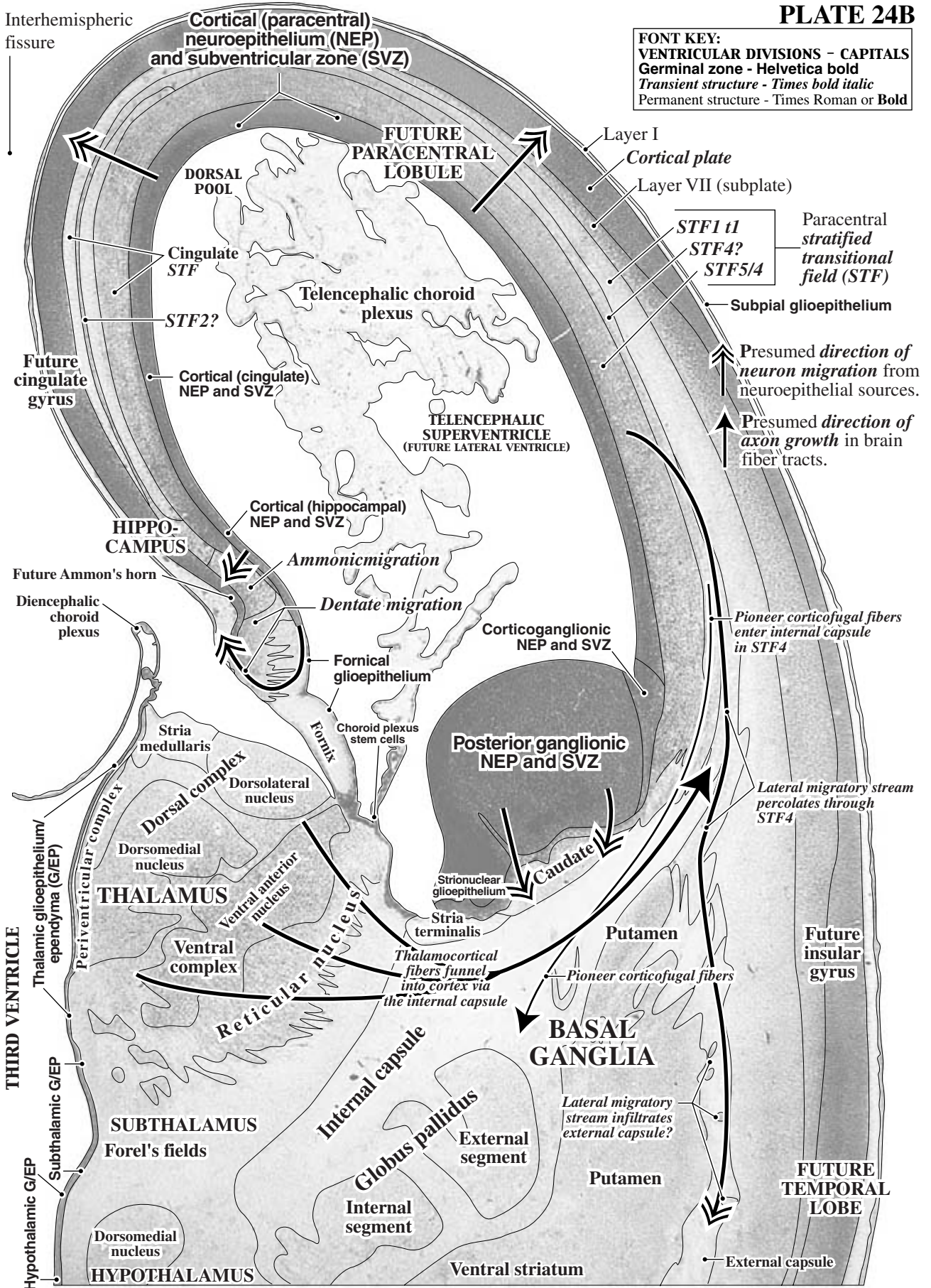
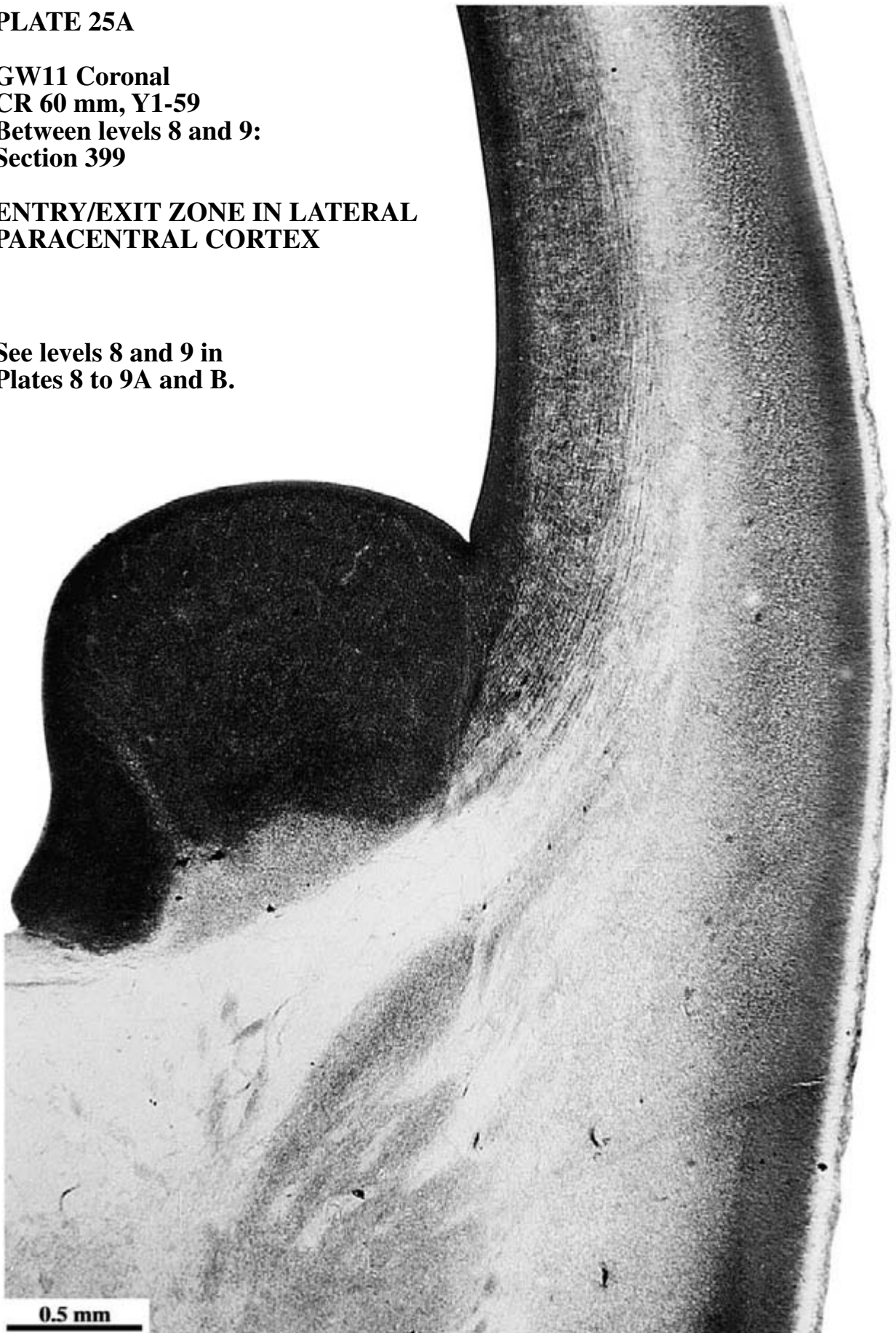


PLATE 25A

**GW11 Coronal
CR 60 mm, Y1-59
Between levels 8 and 9:
Section 399**

**ENTRY/EXIT ZONE IN LATERAL
PARACENTRAL CORTEX**

**See levels 8 and 9 in
Plates 8 to 9A and B.**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

TELENCEPHALIC SUPERVENTRICLE
 (FUTURE LATERAL VENTRICLE)
DORSAL POOL

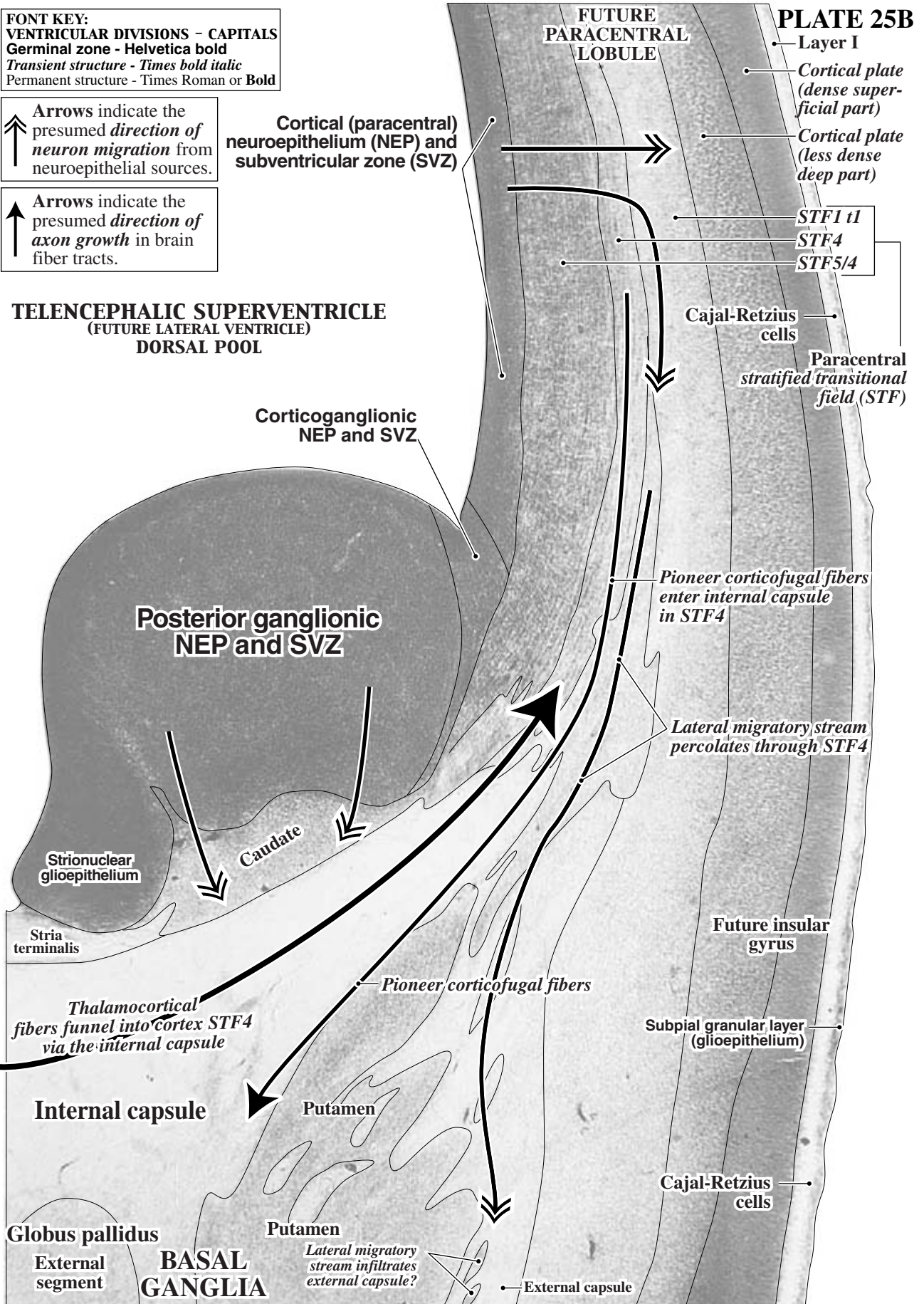


PLATE 26A
GW11 Coronal, CR 60 mm, Y1-59, Near level 8: Section 389

DIENCEPHALON, BASAL GANGLIA,
and BASAL TELENCEPHALON



See level 8 in Plates 8A and B.

PLATE 26B

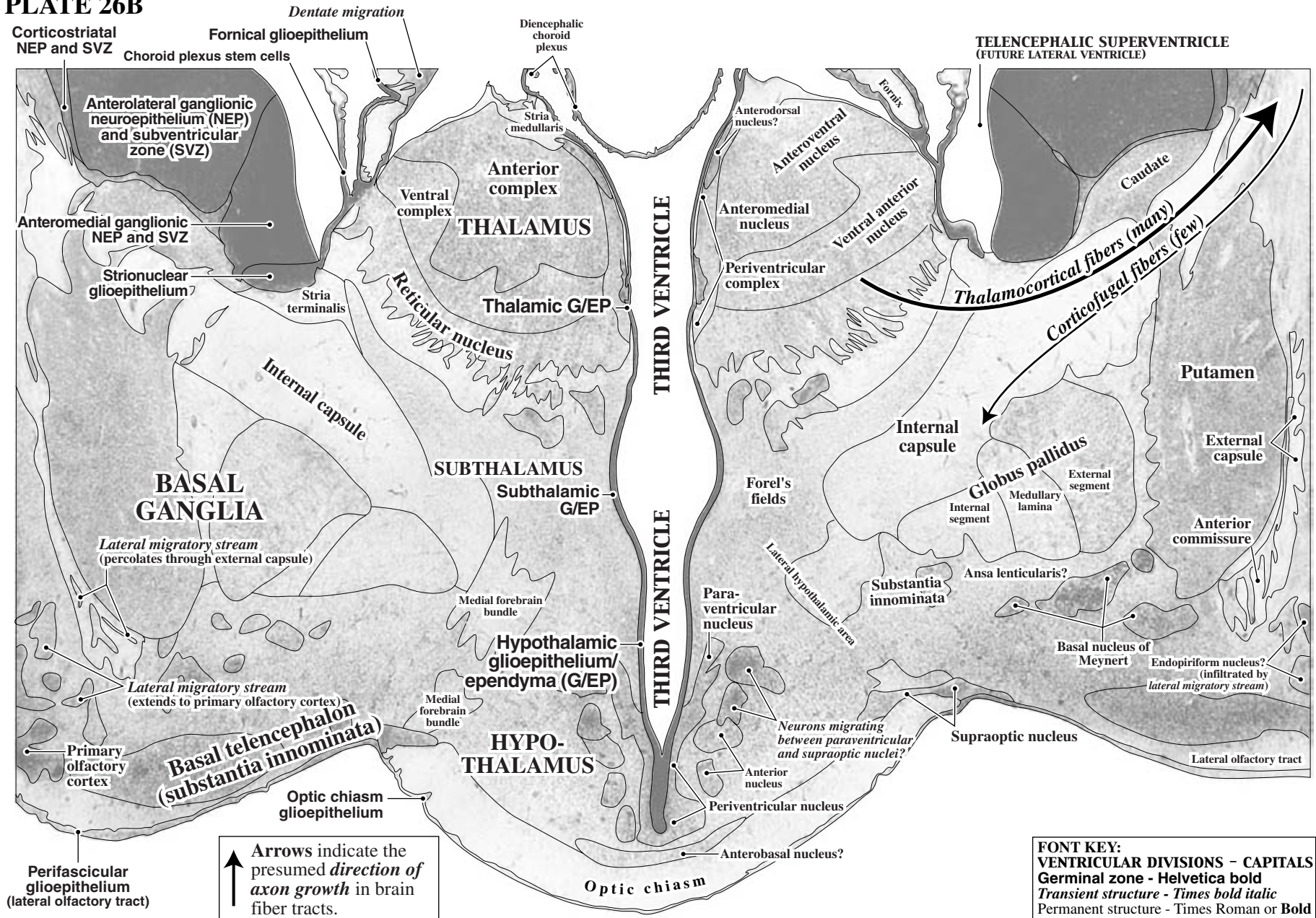


PLATE 27A, GW11 Coronal, CR 60 mm, Y1-59, Level 9: Section 419

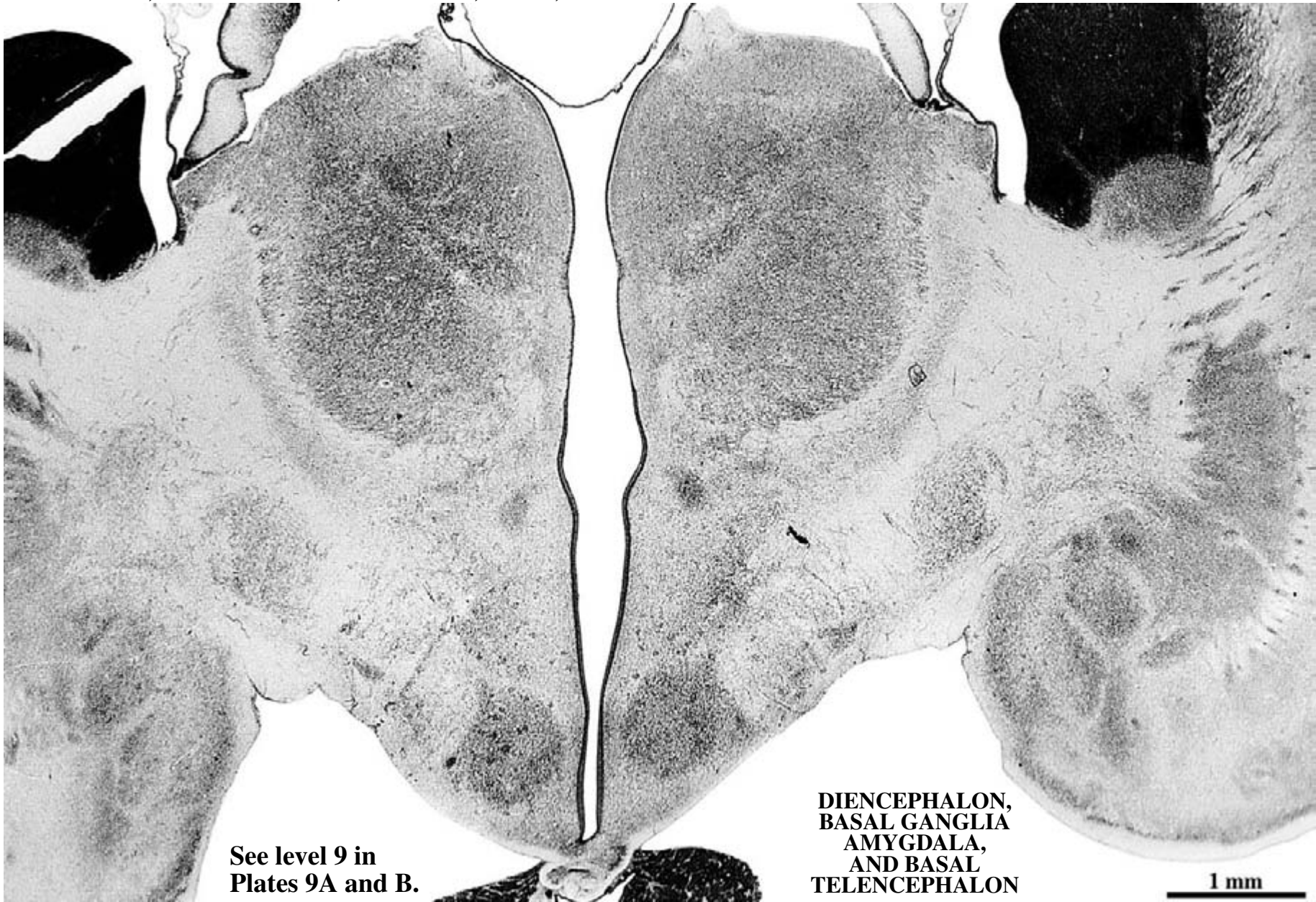


PLATE 27B

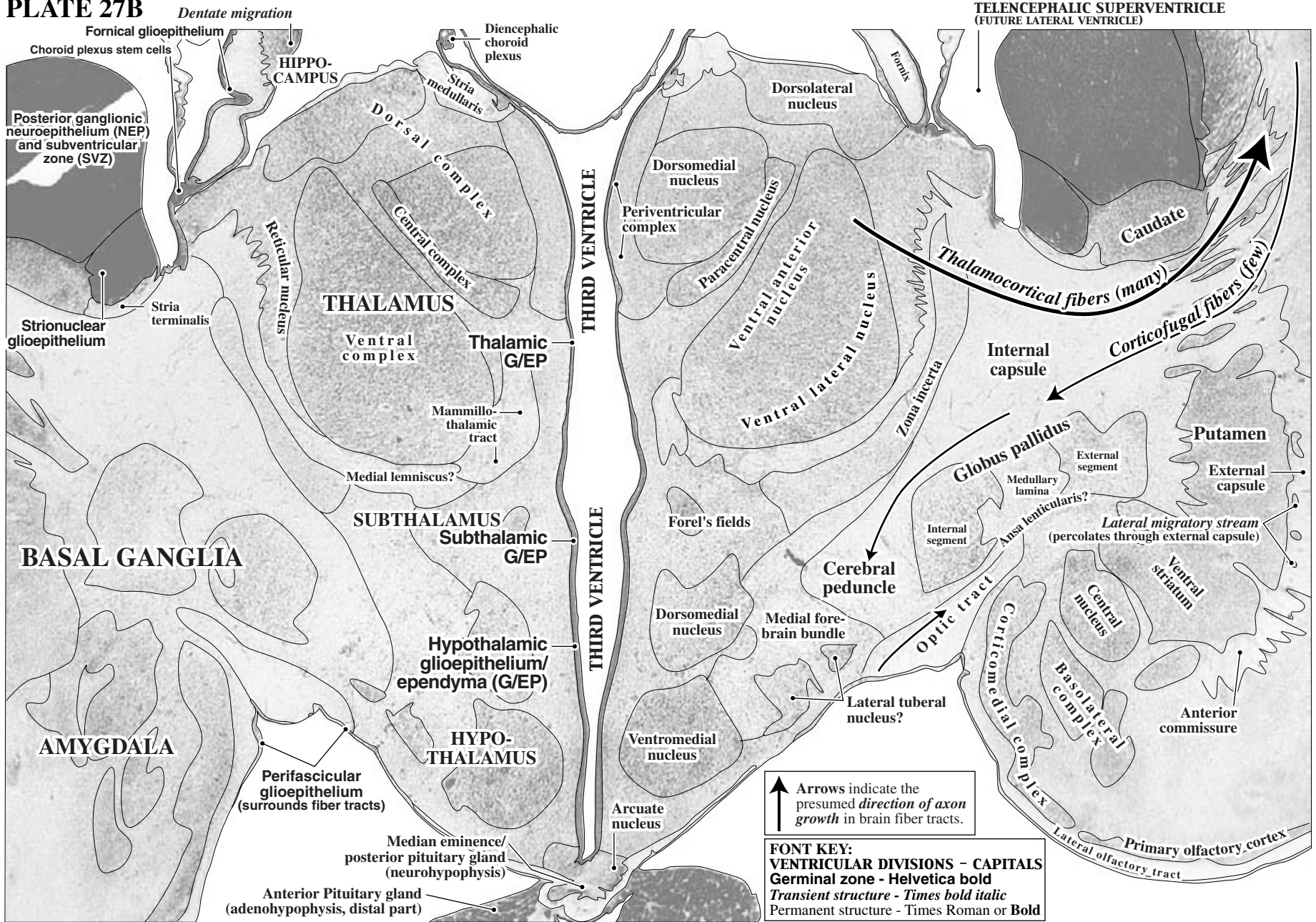
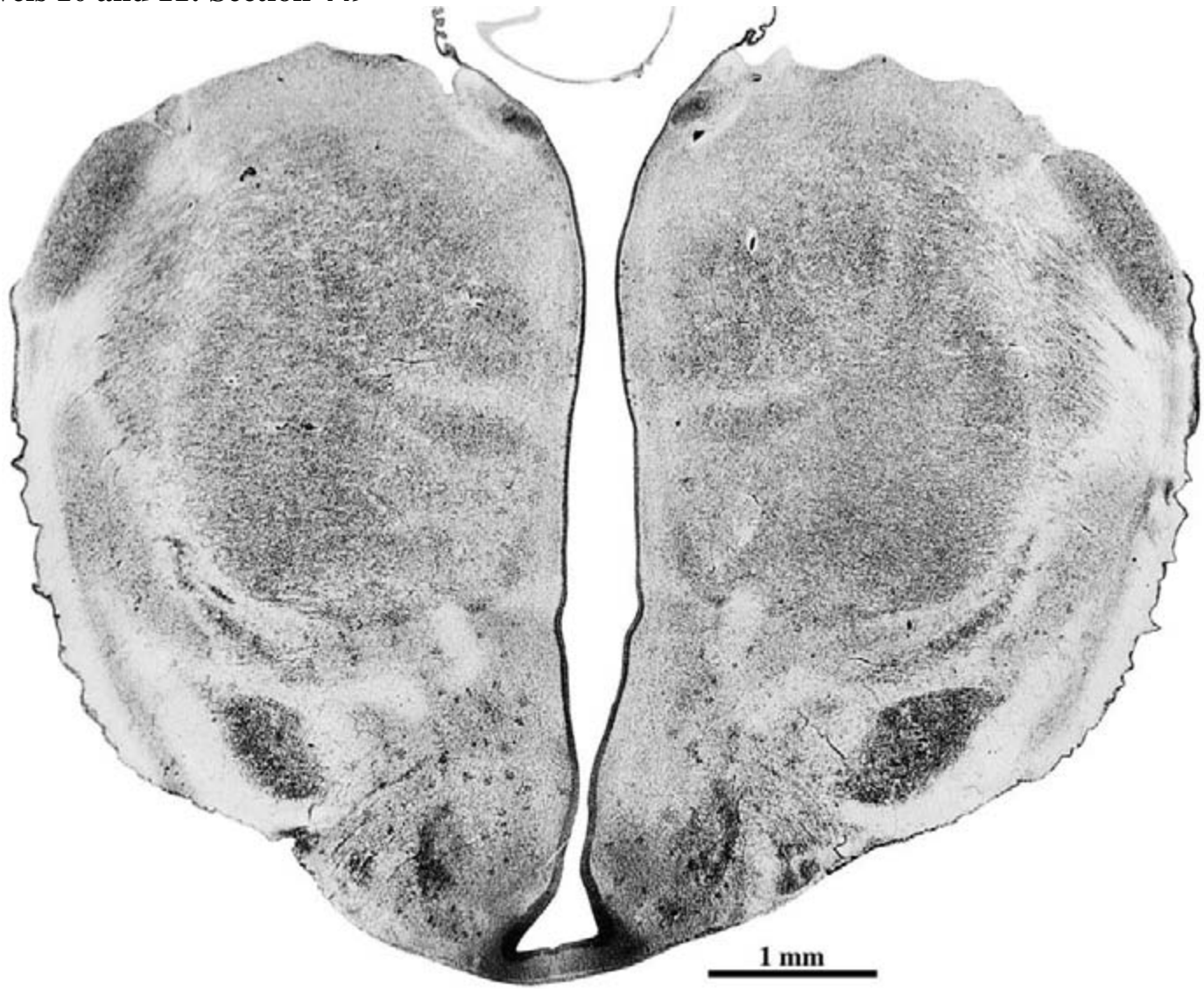


PLATE 28A
GW11 Coronal, CR 60 mm, Y1-59,
Between levels 10 and 11: Section 449

DIENCEPHALON



See levels 10 and 11 in Plates 10A and B to 11A and B.

PLATE 28B

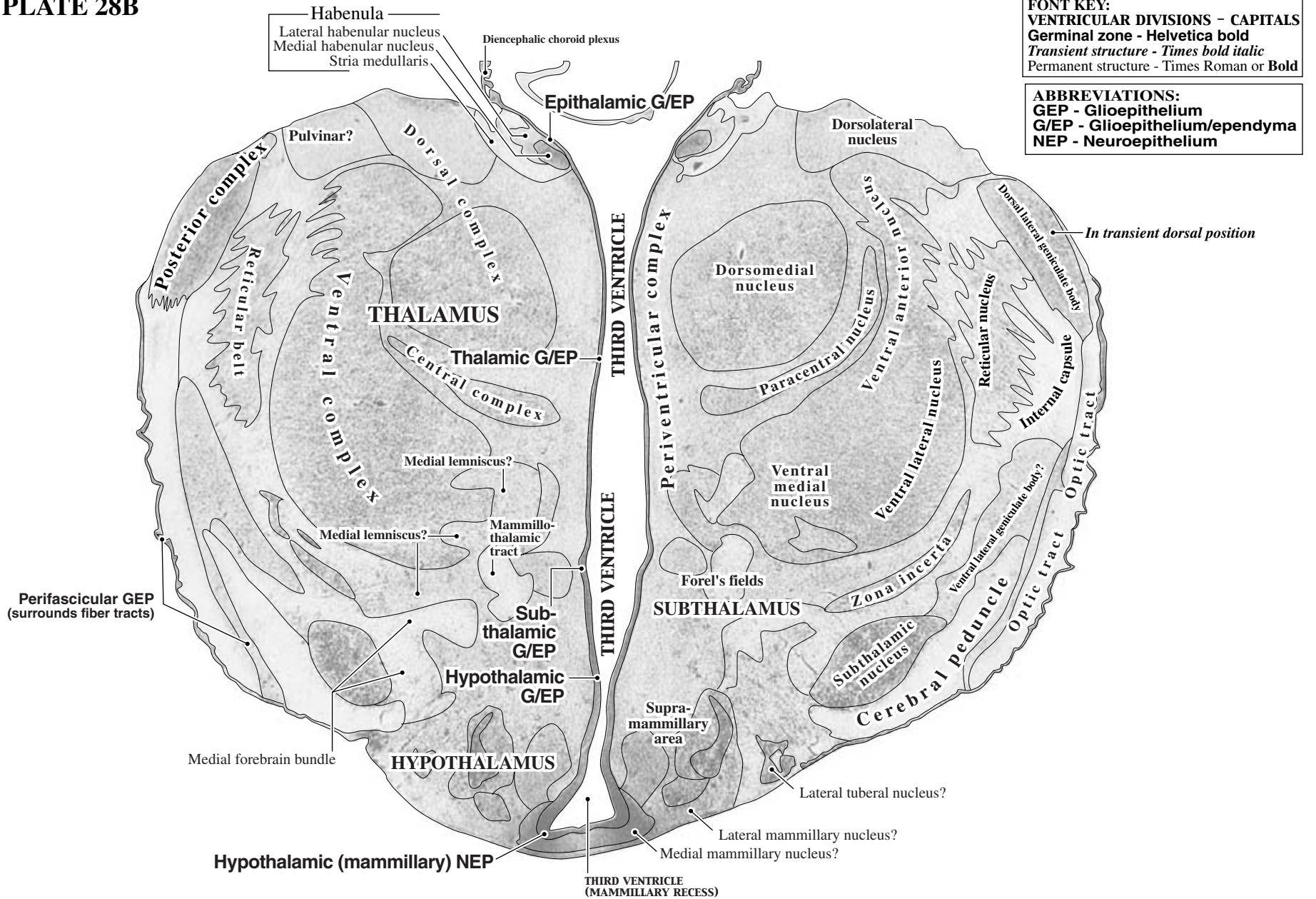
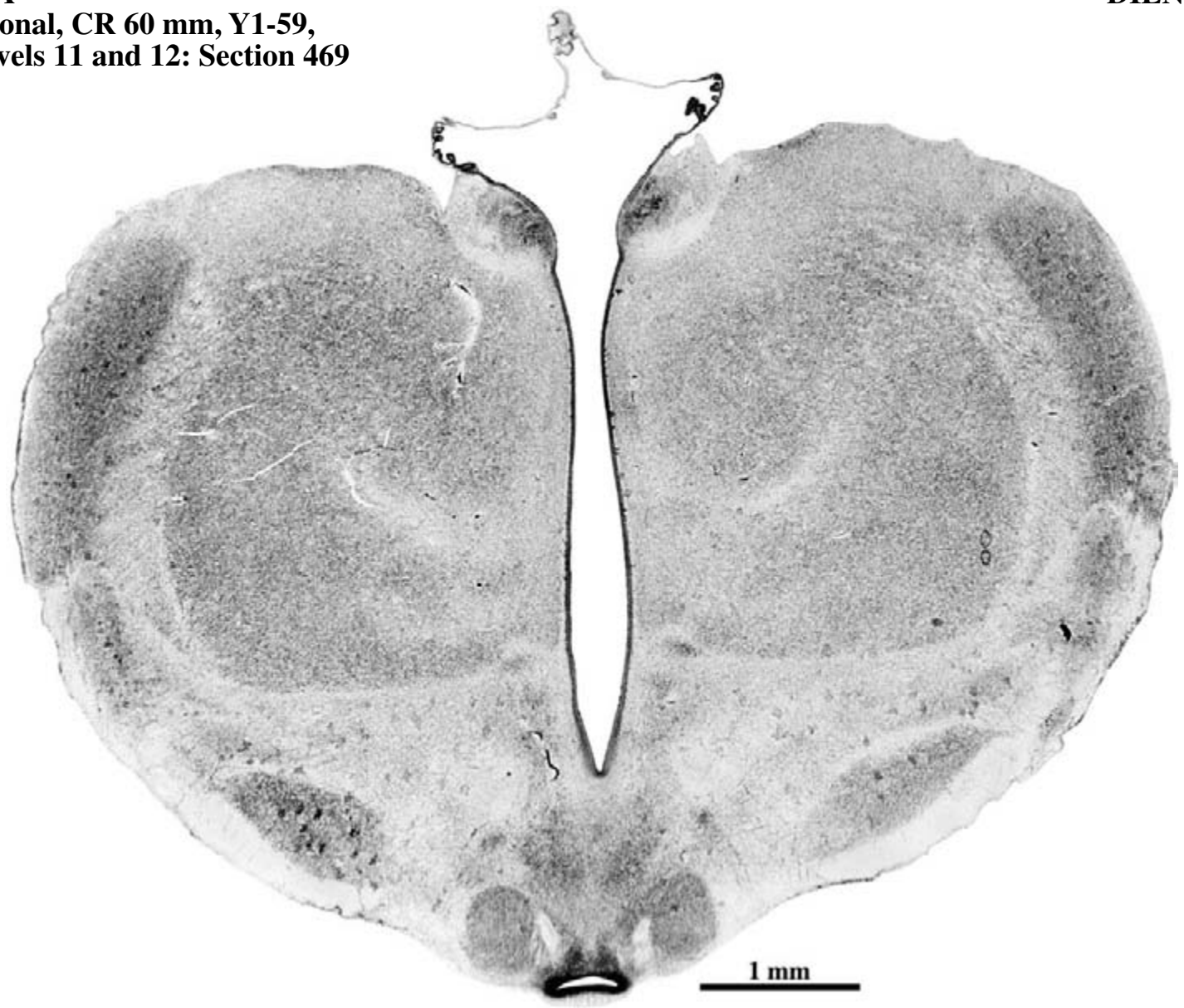


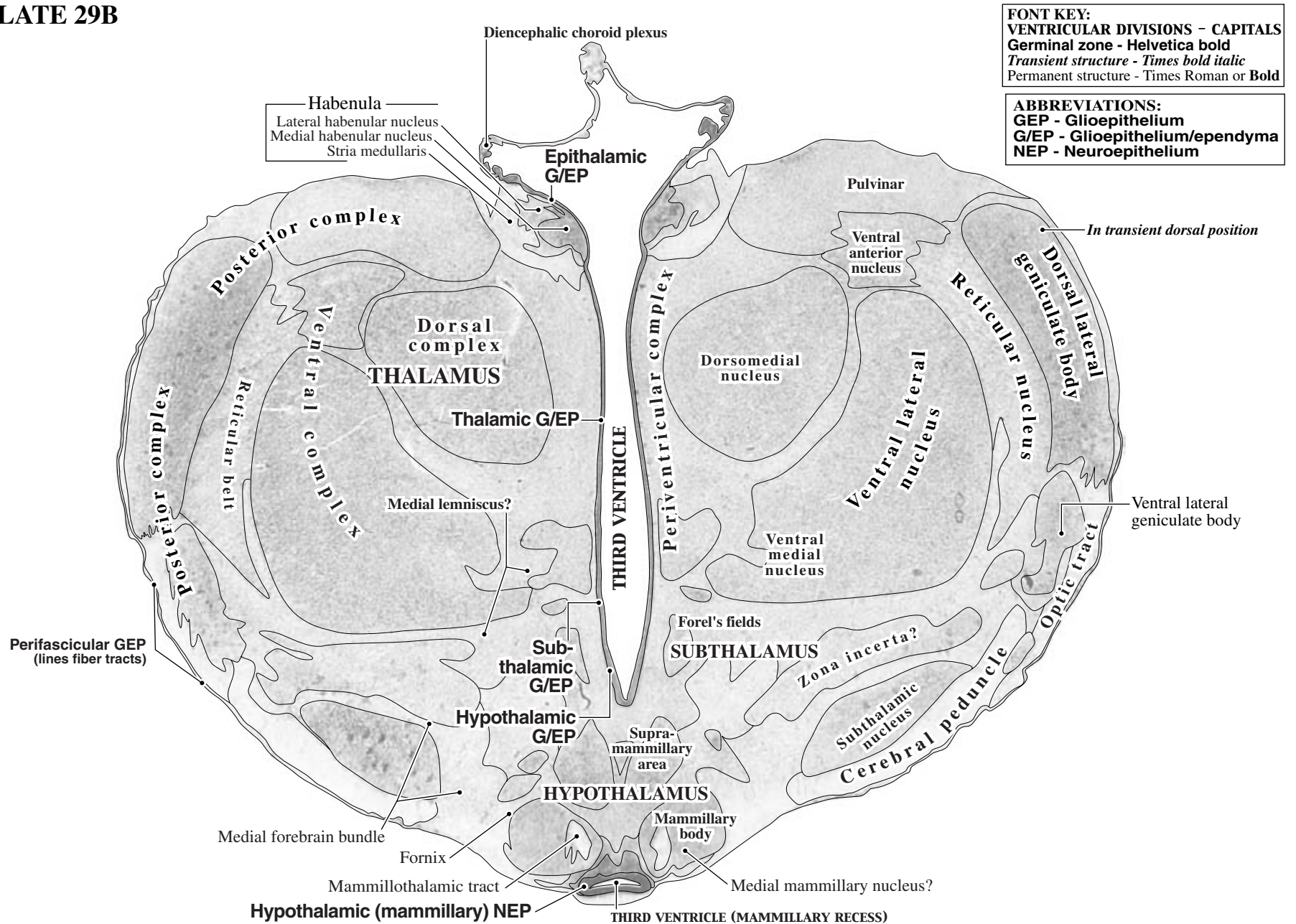
PLATE 29A
GW11 Coronal, CR 60 mm, Y1-59,
Between levels 11 and 12: Section 469

DIENCEPHALON



See levels 11 and 12 in Plates 11A and B to 12A and B.

PLATE 29B

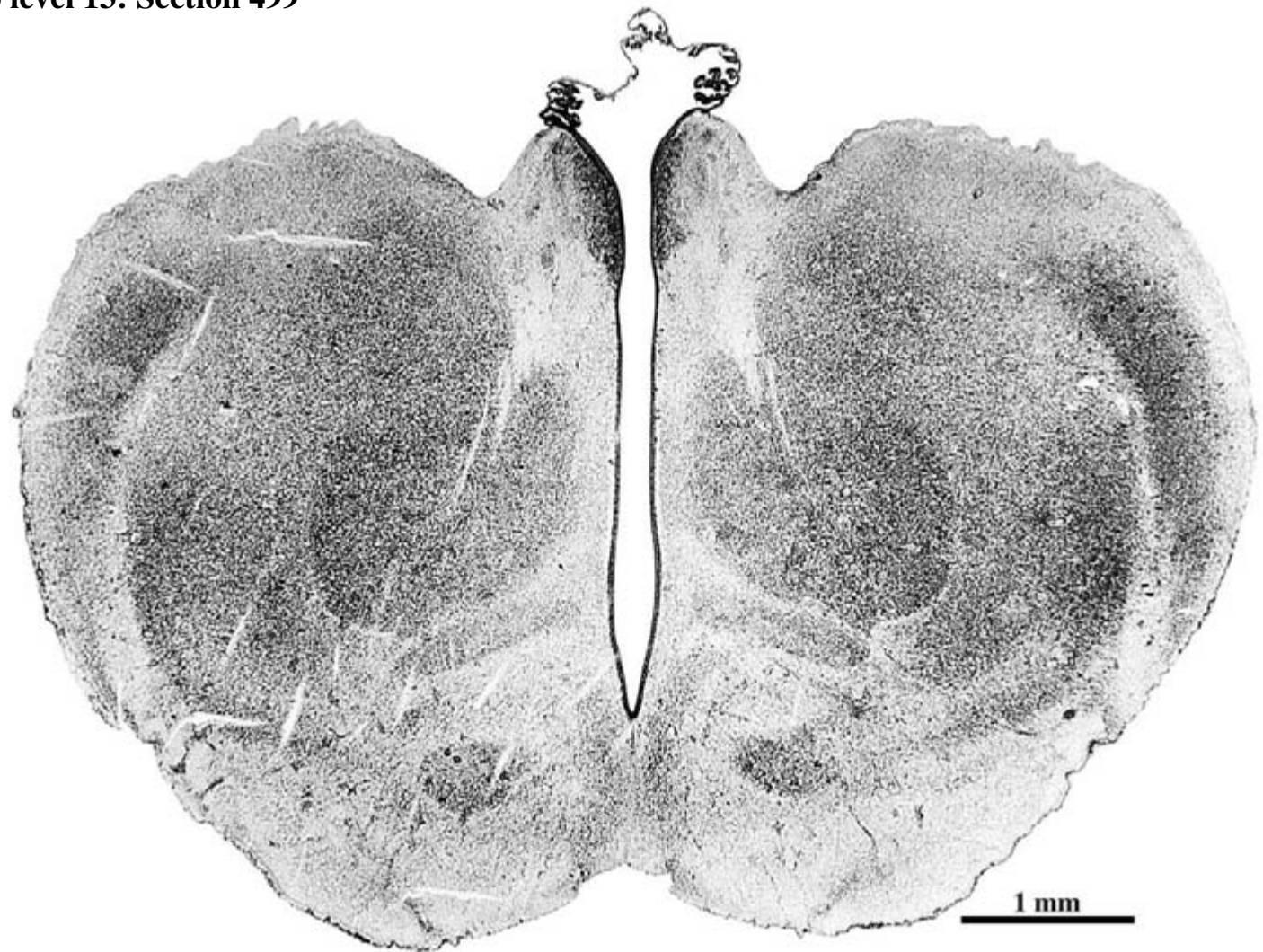


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
G/EP - Glioeptithelium
G/EP - Glioeptithelium/ependyma
NEP - Neuroepithelium

PLATE 30A
GW11 Coronal, CR 60 mm, Y1-59,
Adjacent to level 13: Section 499

DIENCEPHALON



See level 13 in Plates 13A and B.

PLATE 30B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioeptihelium
G/EP - Glioeptihelium/ependyma

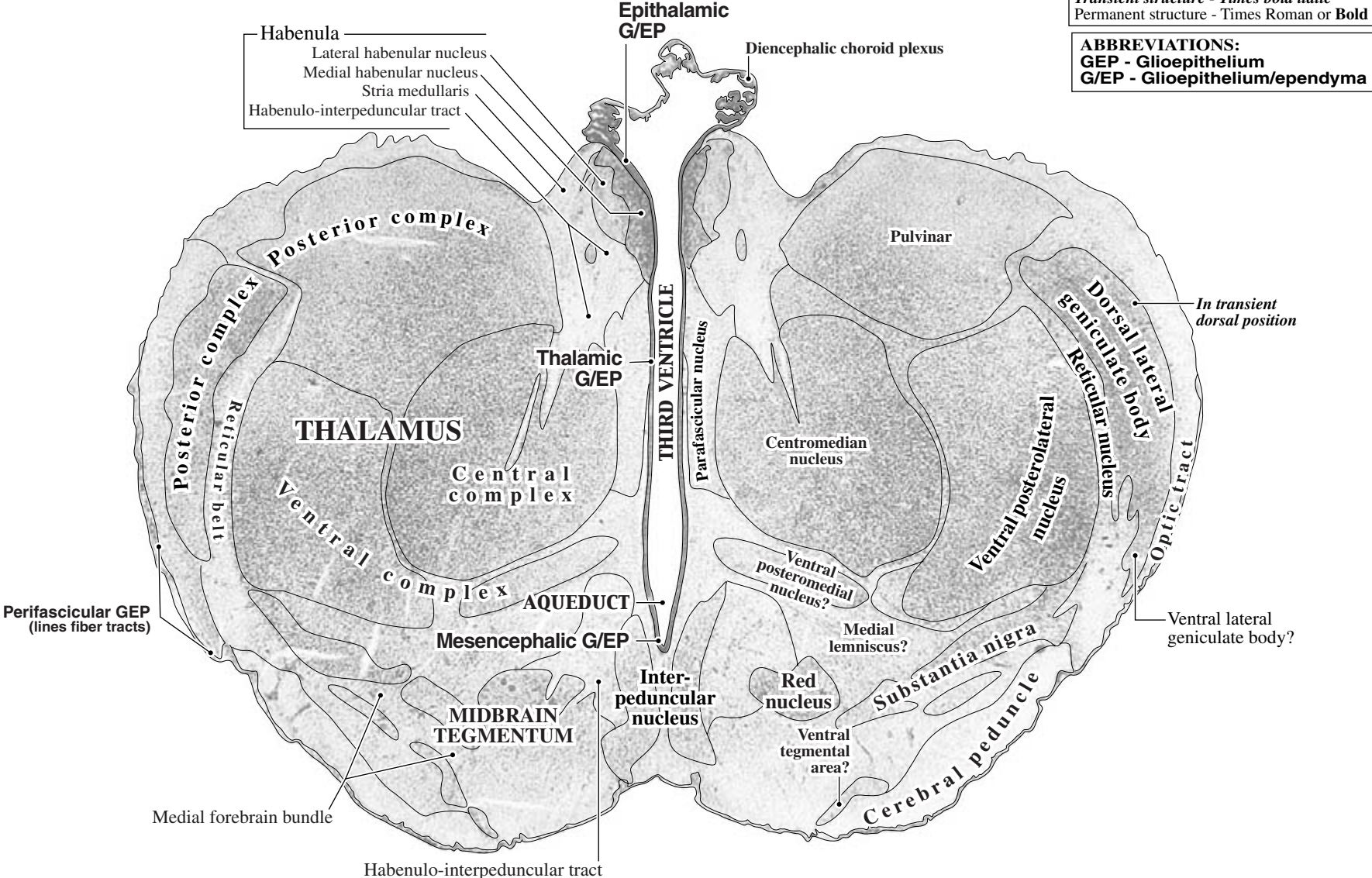
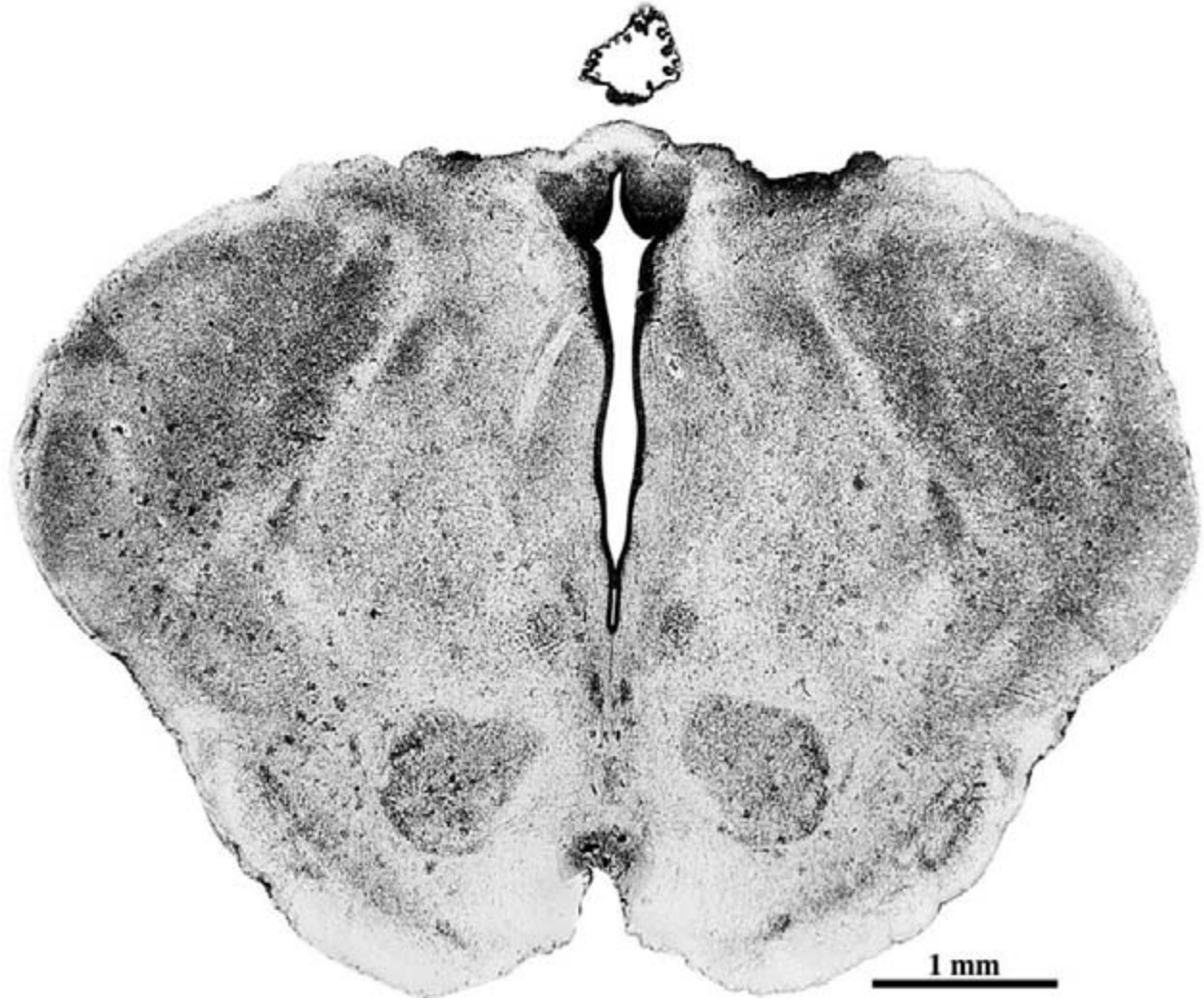


PLATE 31A
GW11 Coronal, CR 60 mm, Y1-59,
Level 14: Section 529

DIENCEPHALON AND MIDBRAIN TEGMENTUM



See level 14 in Plates 14A and B.

PLATE 31B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
G/EP - Glioepithelium
G/EP - Glioepithelium/ependyma

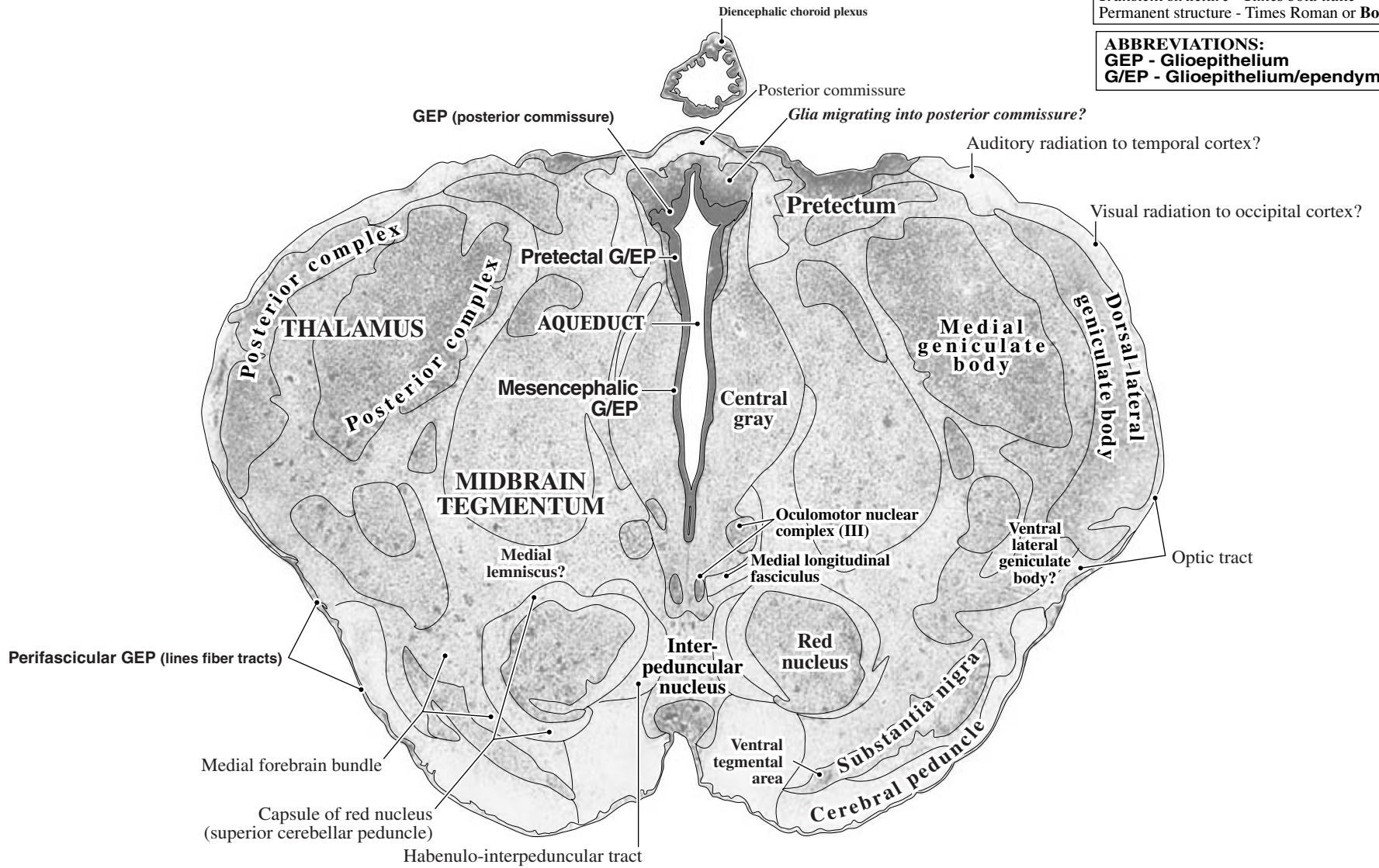
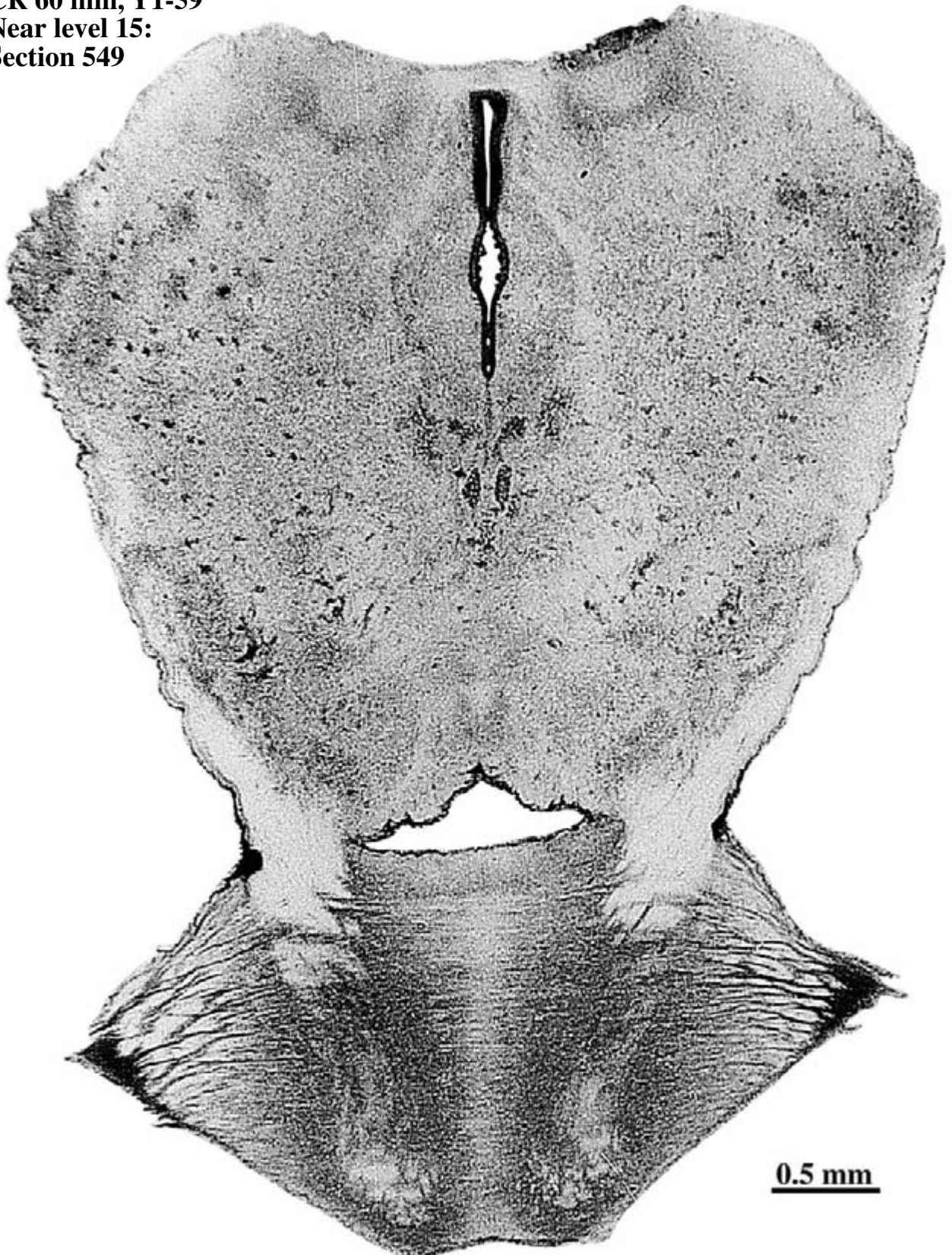


PLATE 32A

MIDBRAIN AND PONS

GW11 Coronal
CR 60 mm, Y1-59
Near level 15:
Section 549



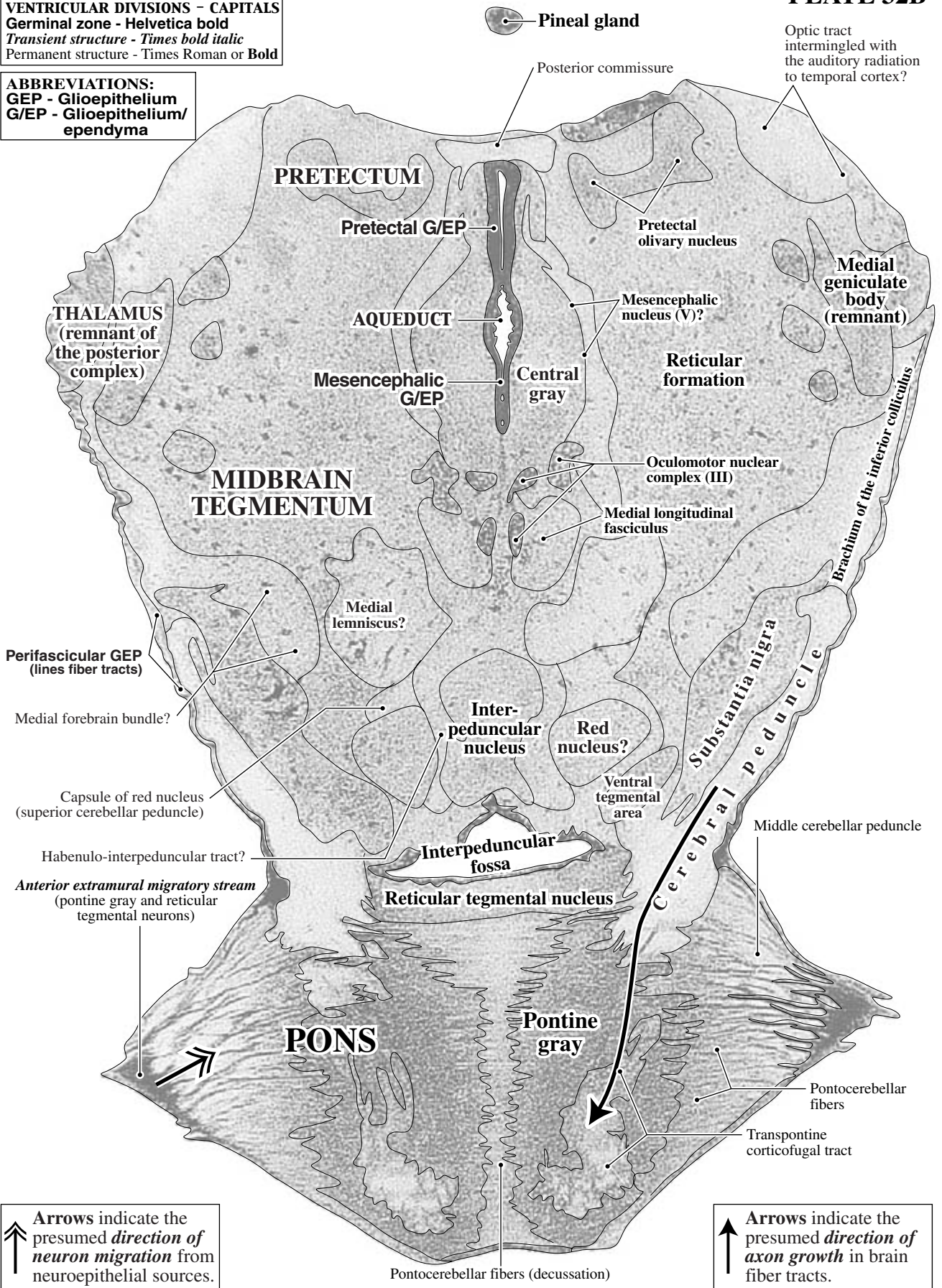
0.5 mm

See Level 15 in Plates 15A and B.

PLATE 32B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma

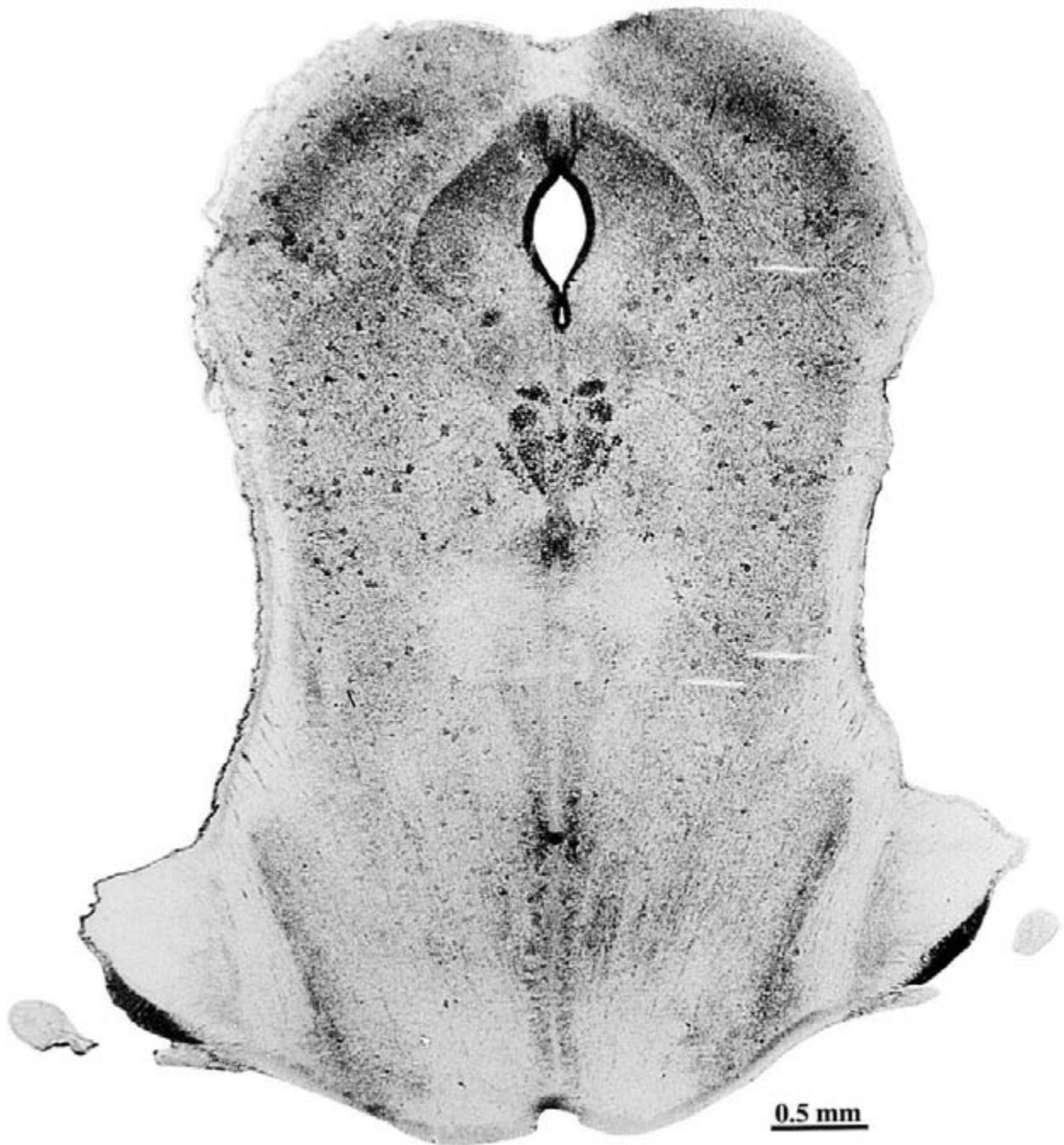


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 33A**MIDBRAIN AND PONS**

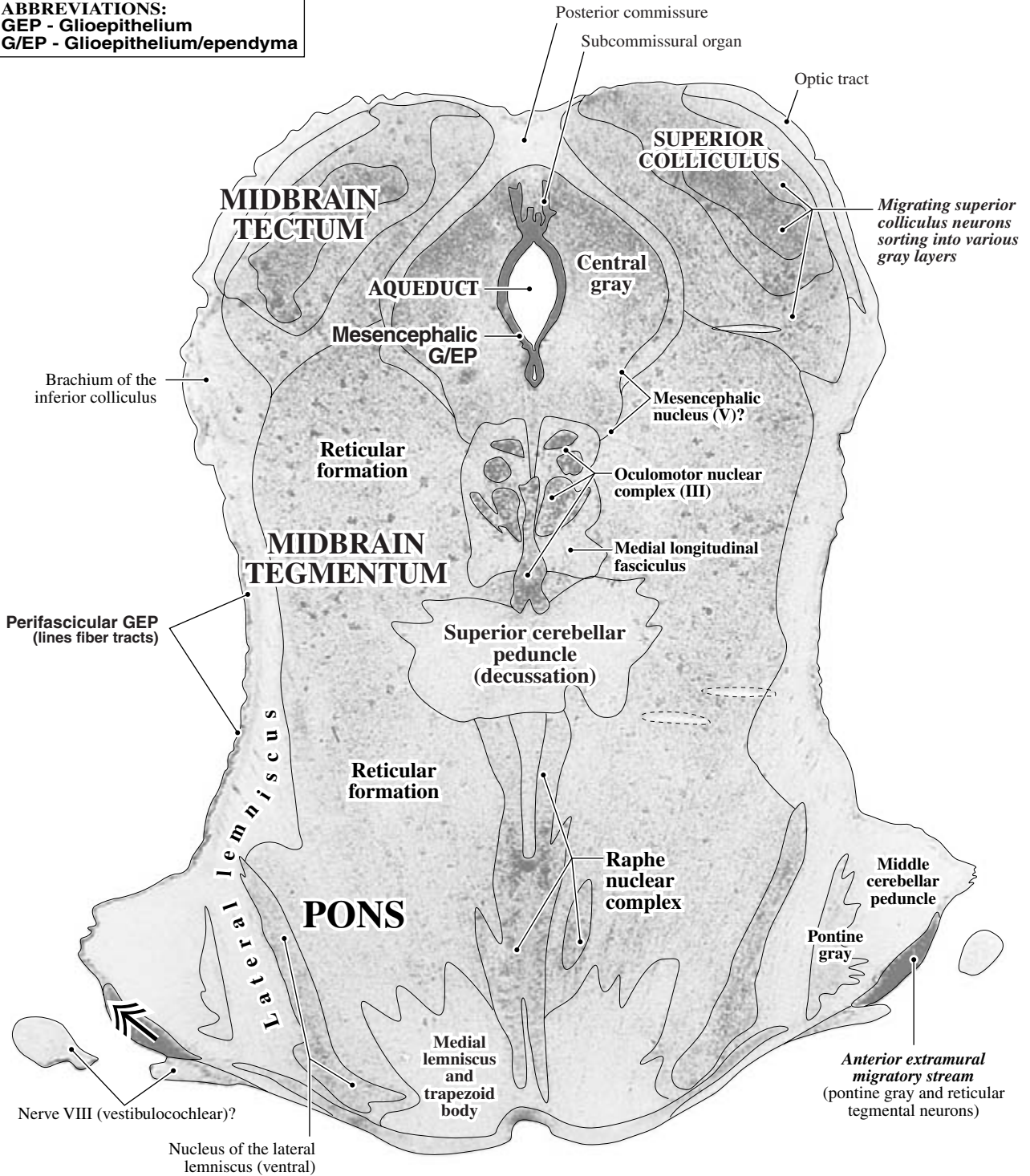
**GW11 Coronal
CR 60 mm, Y1-59
Level 15:
Section 569**



See Level 15 in Plates 15A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Gliopithelium
G/EP - Gliopithelium/ependyma



Dashed lines indicate staining and/or sectioning artifacts.

Arrows indicate the presumed **direction of neuron migration** from neuroepithelial sources.

PLATE 34A**MIDBRAIN, PONS, AND MEDULLA**

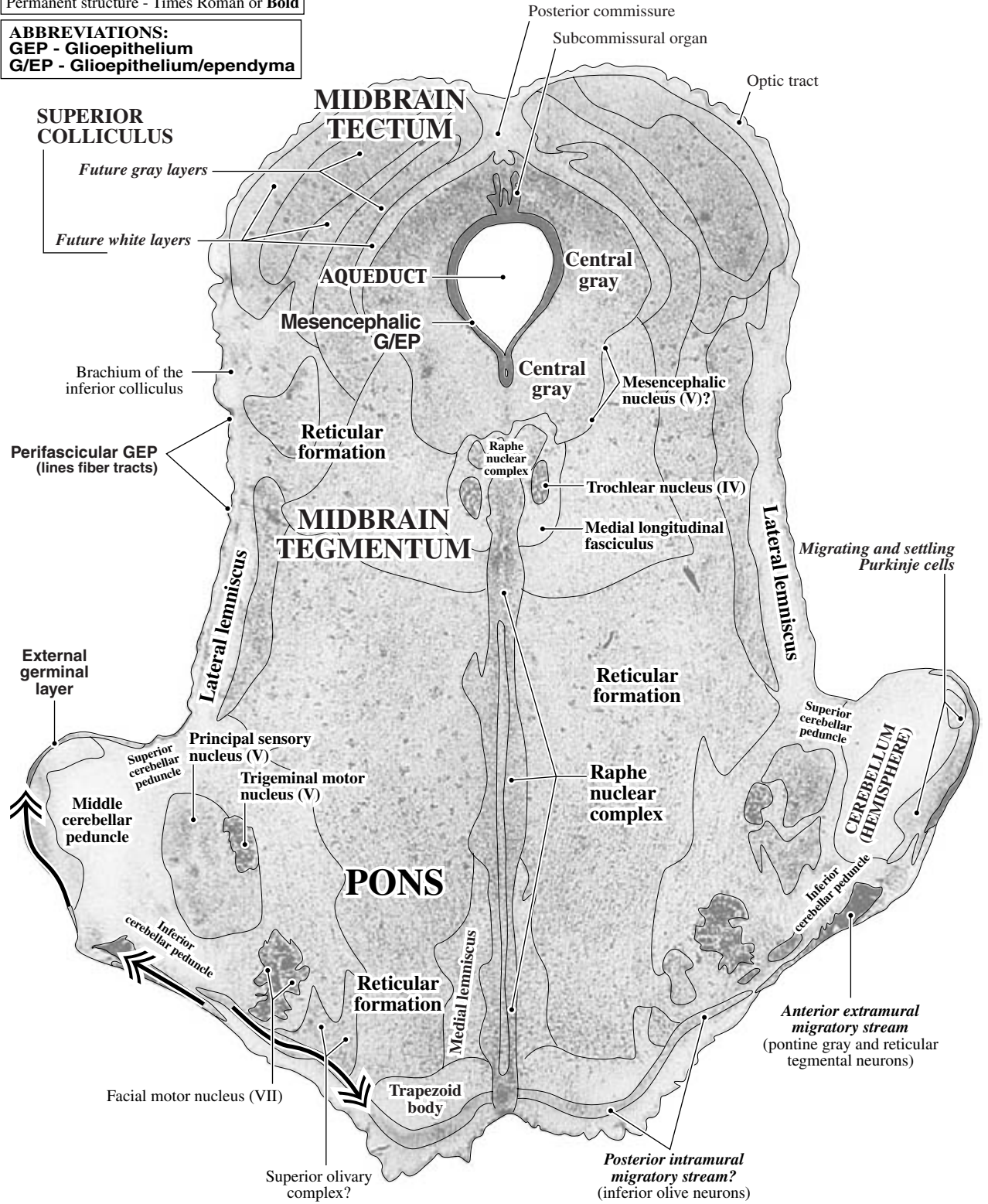
**GW11 Coronal
CR 60 mm, Y1-59
Near level 16:
Section 589**



See Level 16 in Plates 16A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - **Helvetica bold**
 Transient structure - *Times bold italic*
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma



Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 35A**MIDBRAIN, PONS, AND MEDULLA**

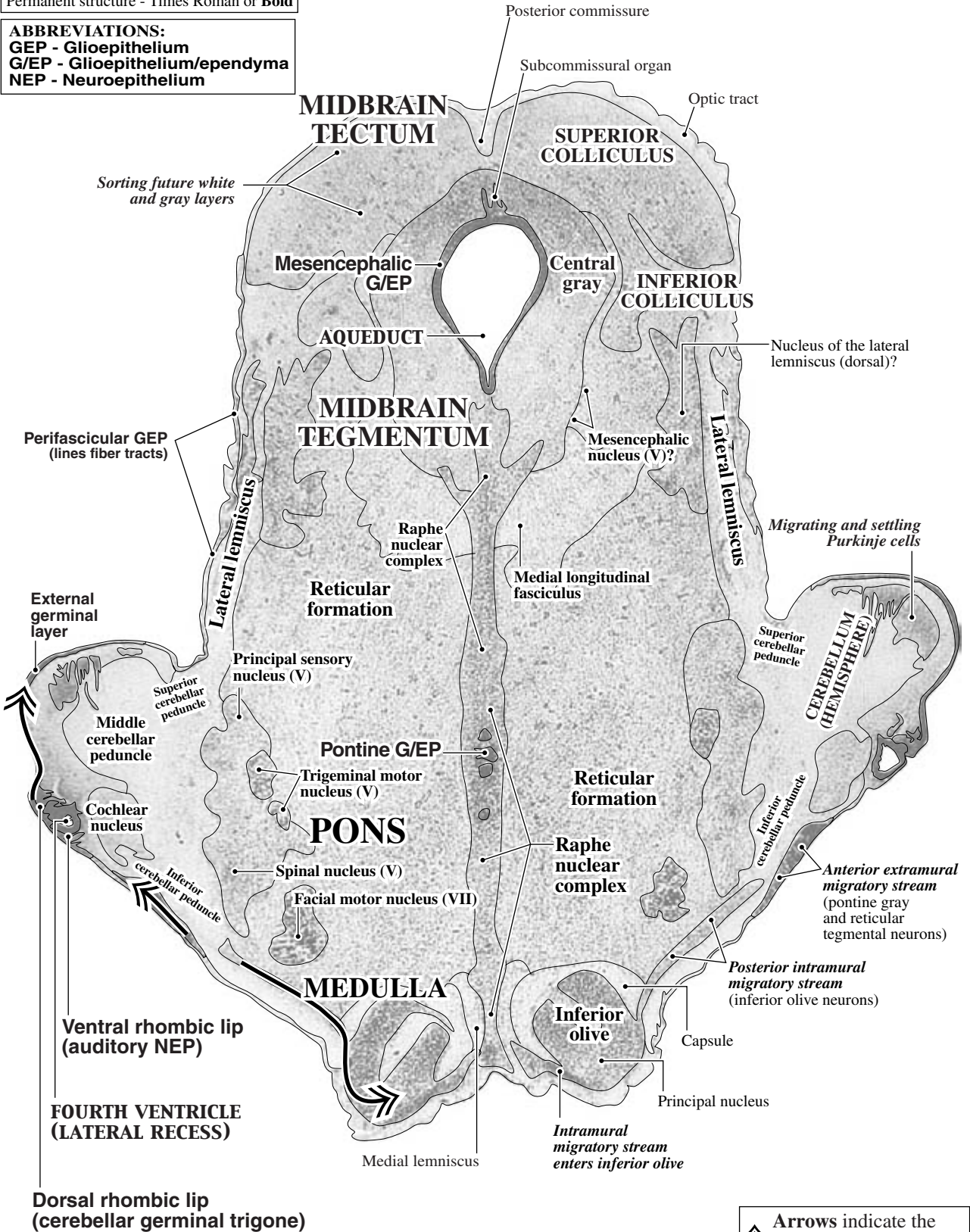
**GW11 Coronal
CR 60 mm, Y1-59
Level 16:
Section 599**



See Level 16 in Plates 16A and B.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
 GEP - Glioepithelium
 G/EP - Glioepithelium/ependyma
 NEP - Neuroepithelium



Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PART III: GW11 HORIZONTAL

This is specimen number 1500 in the Carnegie Collection, designated here as C1500. A normal fetus with a crown-rump length (CR) of 56 mm was collected in 1916. The fetus is estimated to be early in gestational week (GW) 11. The entire fetus was fixed in formalin and was cut in the horizontal plane. The records are not clear regarding section thickness, but the brain is present in 2,400 sections, more than three times the number of 35 μm sections in the GW17 horizontally sectioned brain in the Yakovlev Collection (Bayer and Altman, 2005, Volume 3, Part IX). We estimate that the sections are between 8-10 μm thick. All sections were stained with Bodian's method to show developing fiber tracts. Since there is no photograph of C1500's brain before it was embedded and cut, a specimen from Hochstetter (1919) that is comparable in age is used to show the approximate section plane and external features in early GW11 (**Figure 2**). **Levels 1-10**, large sections containing the cerebral hemispheres, are shown at low magnification in **Plates 36-45**. **Levels 11-21**, small sections containing only the brainstem, are shown at a higher magnification in **Plates 46-56**. To maximize image size within page space, C1500's sections are rotated 90° (landscape orientation). The anterior part of each section is on the left (page bottom), and the posterior part of each section is on the right (page top).

C1500 has many of the same features as Y1-59, except that it is slightly less mature. Throughout the cerebral cortex, the **neuroepithelium and subventricular zone** are prominent. The **stratified transitional field (STF)** contains **STF1** and **STF5** throughout; with **STF4** only in lateral areas. The most prominent developmental feature of the cerebral cortex is that both the **STF** layers and the cortical plate have a pronounced lateral (thicker) to medial (thinner) maturation gradient. The olfactory bulb beneath the anterior septum and striatum contains a small **rostral migratory stream** in its core. In anterolateral parts of the cerebral cortex, streams of neurons and glia appear to leave **STF4** and enter the **lateral migratory stream**. The hippocampus is in an immature position dorsal to the thalamus and medial to the temporal lobe. Cells are entering Ammon's horn pyramidal layer in the **ammonic migration**, and granule cells and their precursors are migrating to the hilus of the presumptive dentate gyrus in the **den-**

tate migration; there is no granular layer. A massive **neuroepithelium/subventricular zone** overlies the amygdala, nucleus accumbens, and striatum (caudate and putamen) where neurons (and glia) are being generated.

The cerebellum is a thick, smooth plate overlying the posterior pons and medulla. However, there is only a thin **glioepithelium/ependyma** at the ventricular surface, indicating that all deep neurons and Purkinje cells have been generated. The deep neurons are in place beneath the cortex, but have indefinite nuclear subdivisions. The cortical surface is covered by an **external germinal layer (egl)** that is actively producing neuronal stem cells, granule, stellate, and basket cells of the cerebellar cortex. Lamination in the cortex is nearly absent, except for a thin molecular layer beneath the **egl**. Nearly all Purkinje cells are migrating, and settling. In contrast to Y1-59, there is no evidence of lobulation in the cerebellar cortex.

The third ventricle, aqueduct, and fourth ventricle are lined by a thin **glioepithelium/ependyma** indicating that neurogenesis in the primary neuroepithelium is complete. In the medulla there are two active germinal sites in anterior and posterior parts of the ventral rhombic lip. 1) The **auditory neuroepithelium** generates cochlear nucleus neurons. 2) A large **precerebellar neuroepithelium** generates precerebellar (mainly pontine gray) neurons.

Neurons throughout the diencephalon, midbrain tegmentum, pons, and medulla are settling. Because C1500 is not Nissl-stained, nuclear divisions are very indistinct. The large **anterior extramural, posterior extramural, and intramural migratory streams** are prominent in the medulla and pons. The Bodian stain clearly shows several fiber tracts and nerves throughout the brainstem. The optic nerve and tract are well defined, along with the medial forebrain bundle. Unlike Y1-59, there is no sure evidence of a cerebral peduncle in the midbrain tegmentum or pontine gray. However, pontine gray fibers cross the midline and a distinct middle cerebellar peduncle is present. There is also a distinct superior and inferior cerebellar peduncle. There is definite staining in the trigeminal nerve and tract, the facial nerve, the abducens nerve, and the glossopharyngeal nerve.

GW11 HORIZONTAL SECTION PLANES

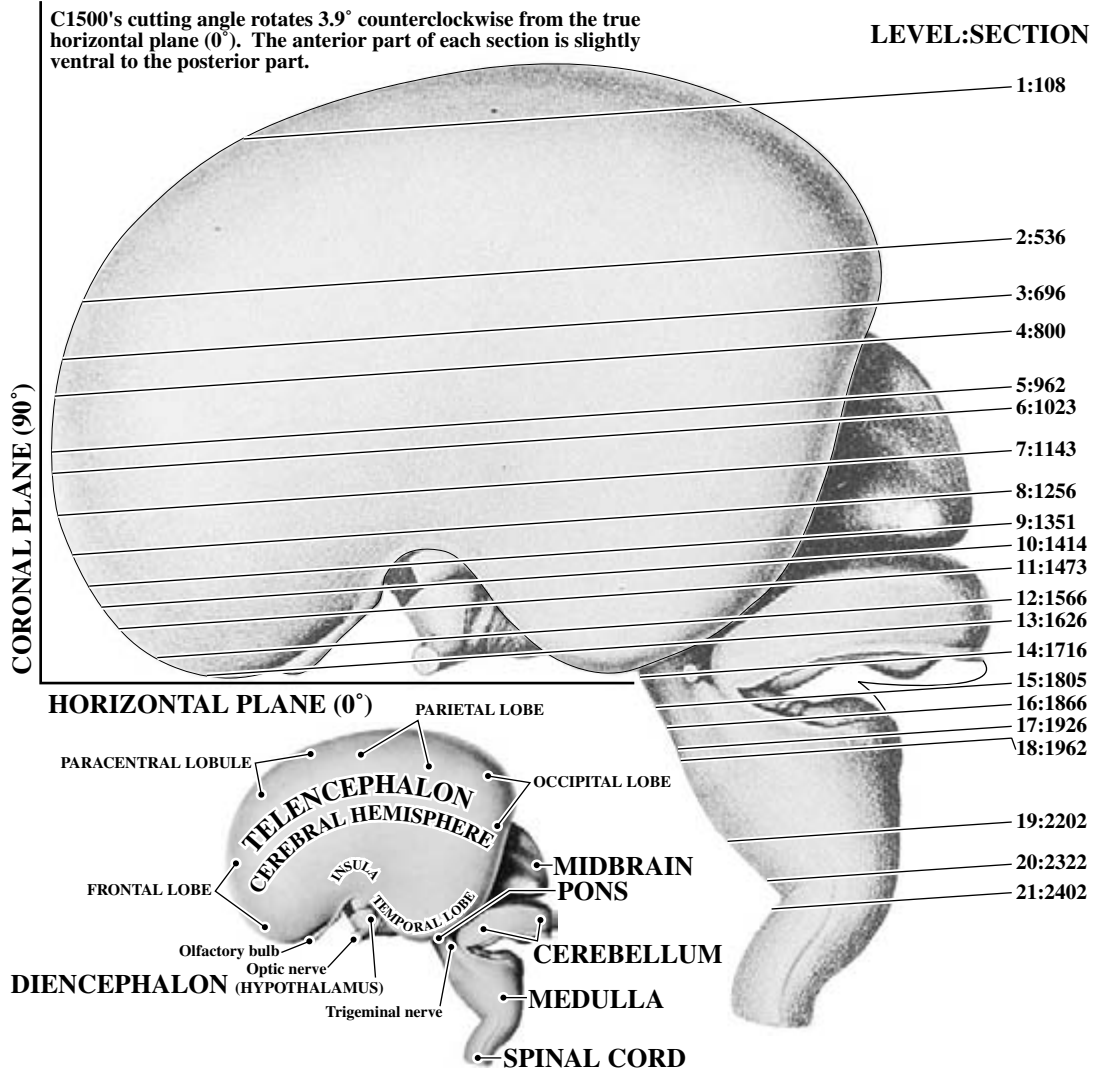


Figure 2. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 53 mm (modified from Figure 46, Table VIII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of C1500 in the following pages. The small inset identifies the major structural features. The cut beneath the cerebellum is the edge of the medullary velum.

PLATE 36A

GW11 Horizontal
CR 57 mm
C1500
Level 1: Section 108

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

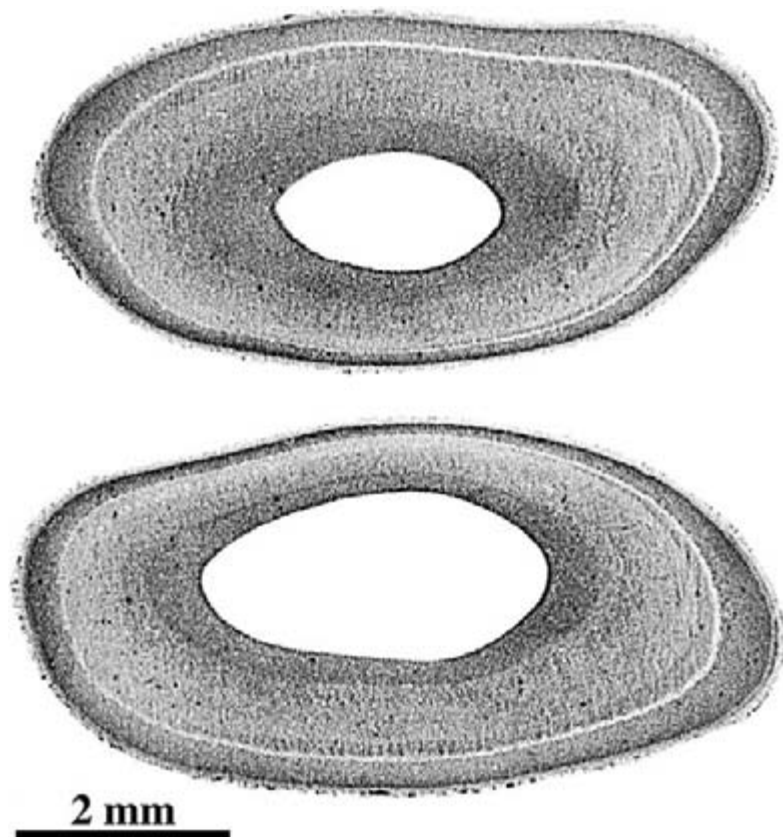


PLATE 36B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

NEP - Neuroepithelium

FUTURE PARACENTRAL LOBULE

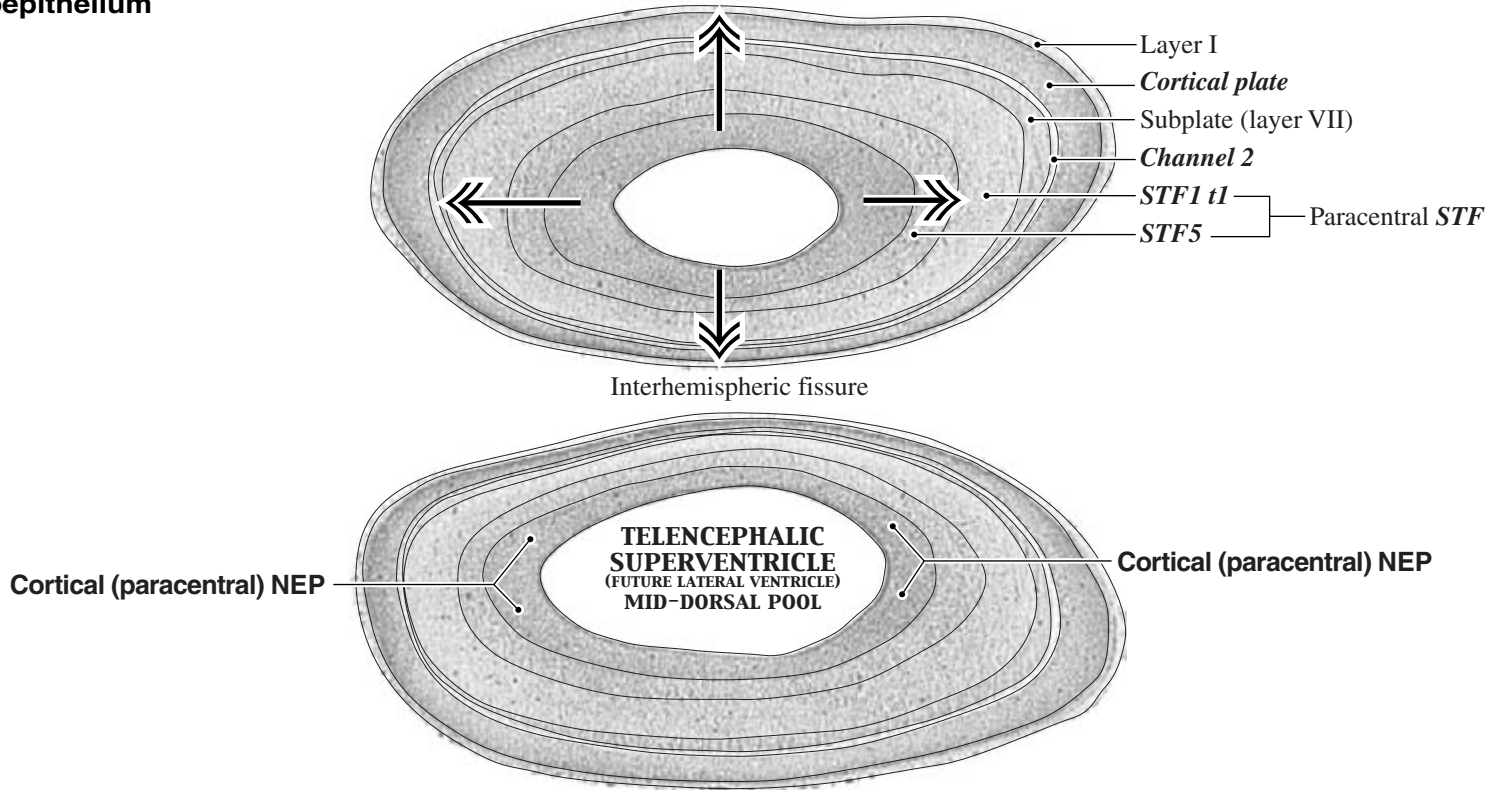
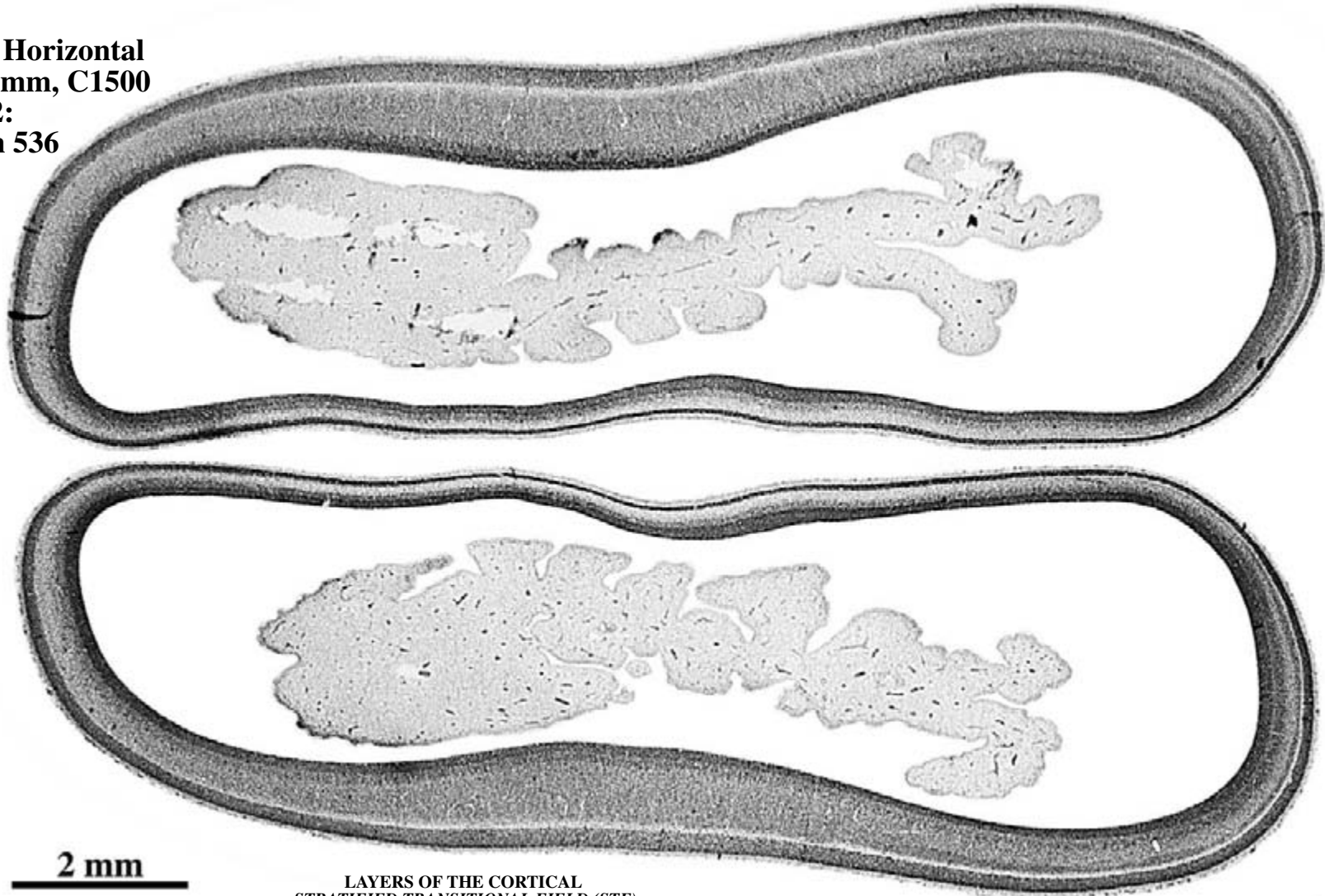


PLATE 37A

**GW11 Horizontal
CR 57 mm, C1500
Level 2:
Section 536**



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 37B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

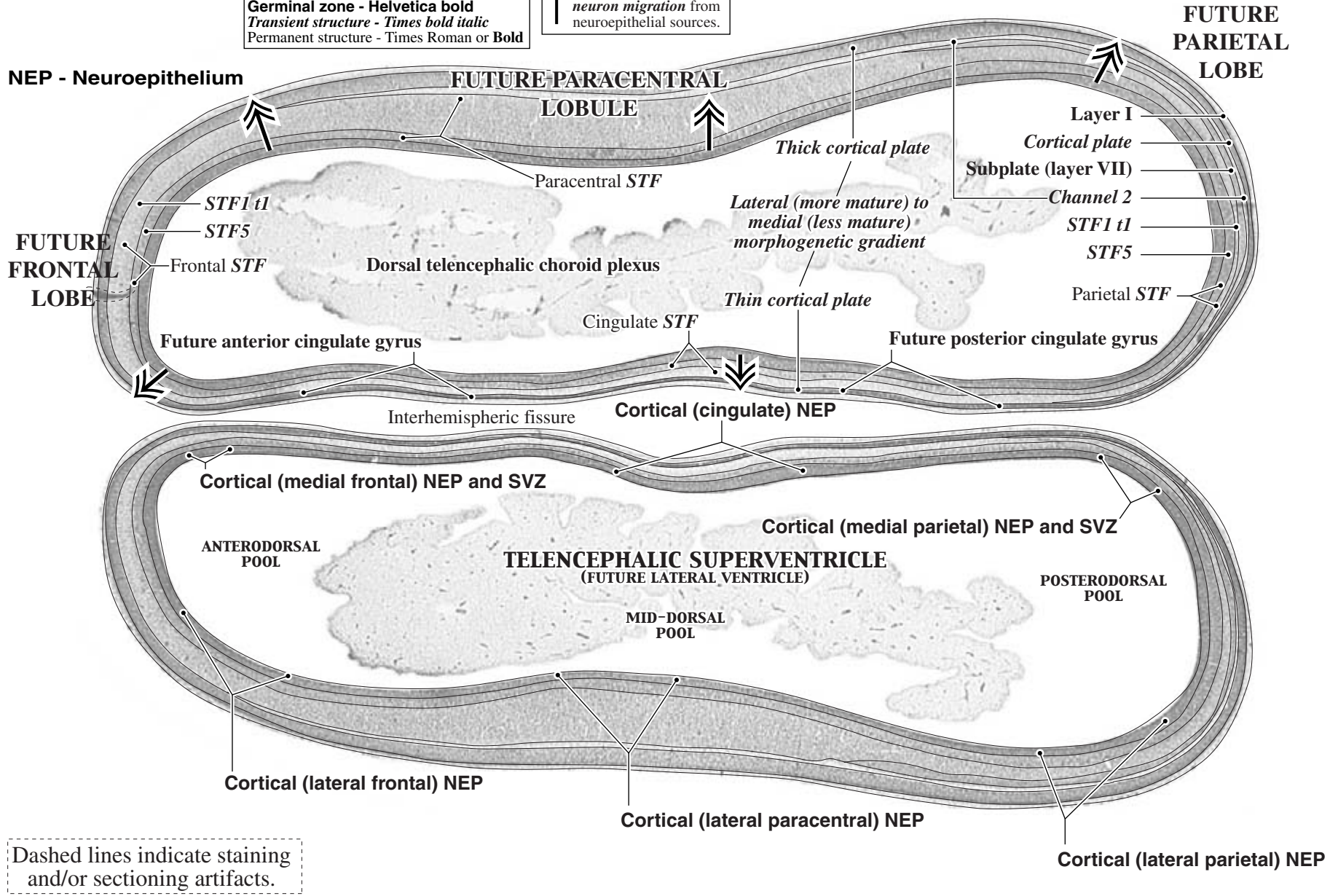
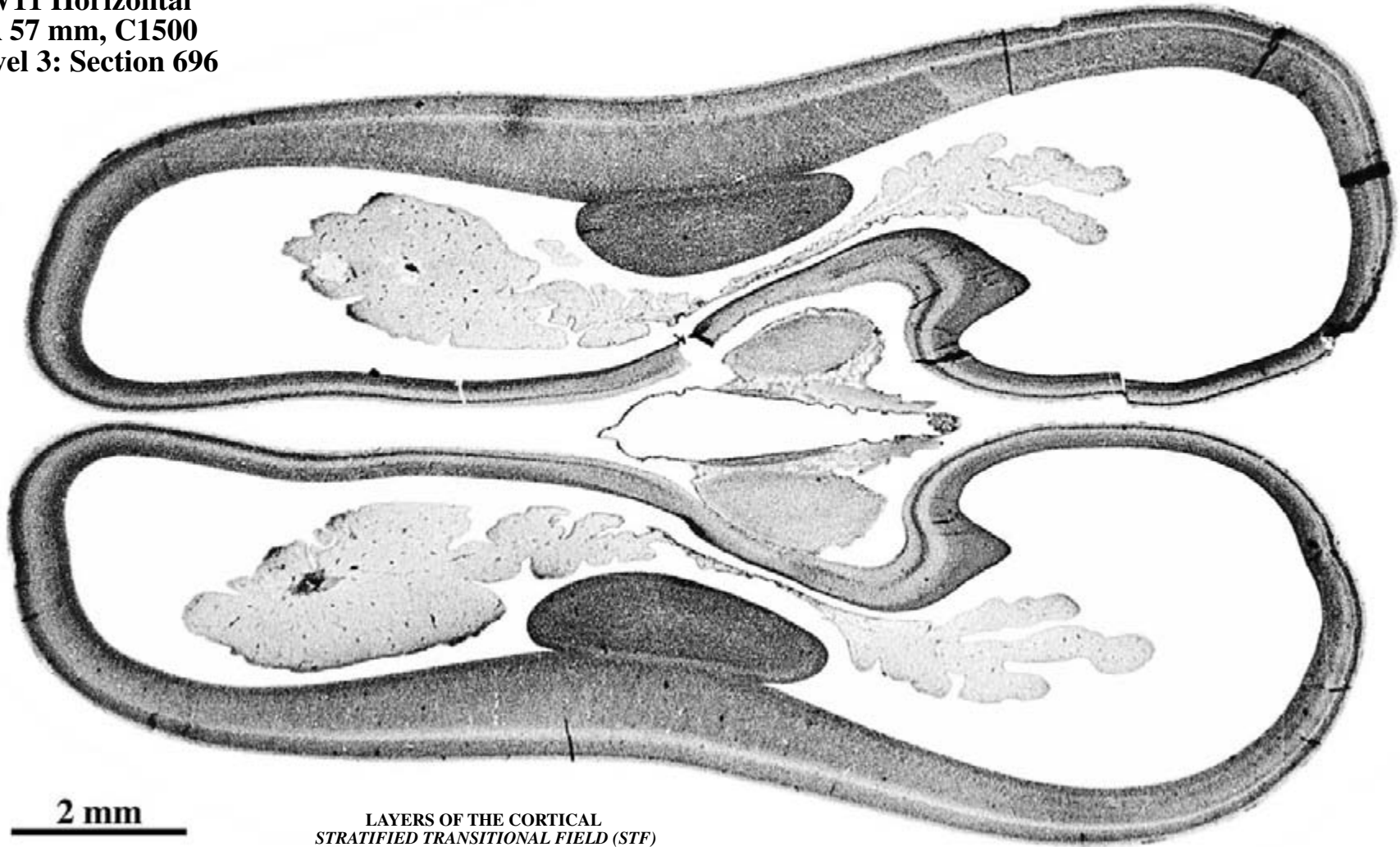


PLATE 38A

**GW11 Horizontal
CR 57 mm, C1500
Level 3: Section 696**



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*I*) when many cells are migrating through it, followed by a late stage (*I2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 38B

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.

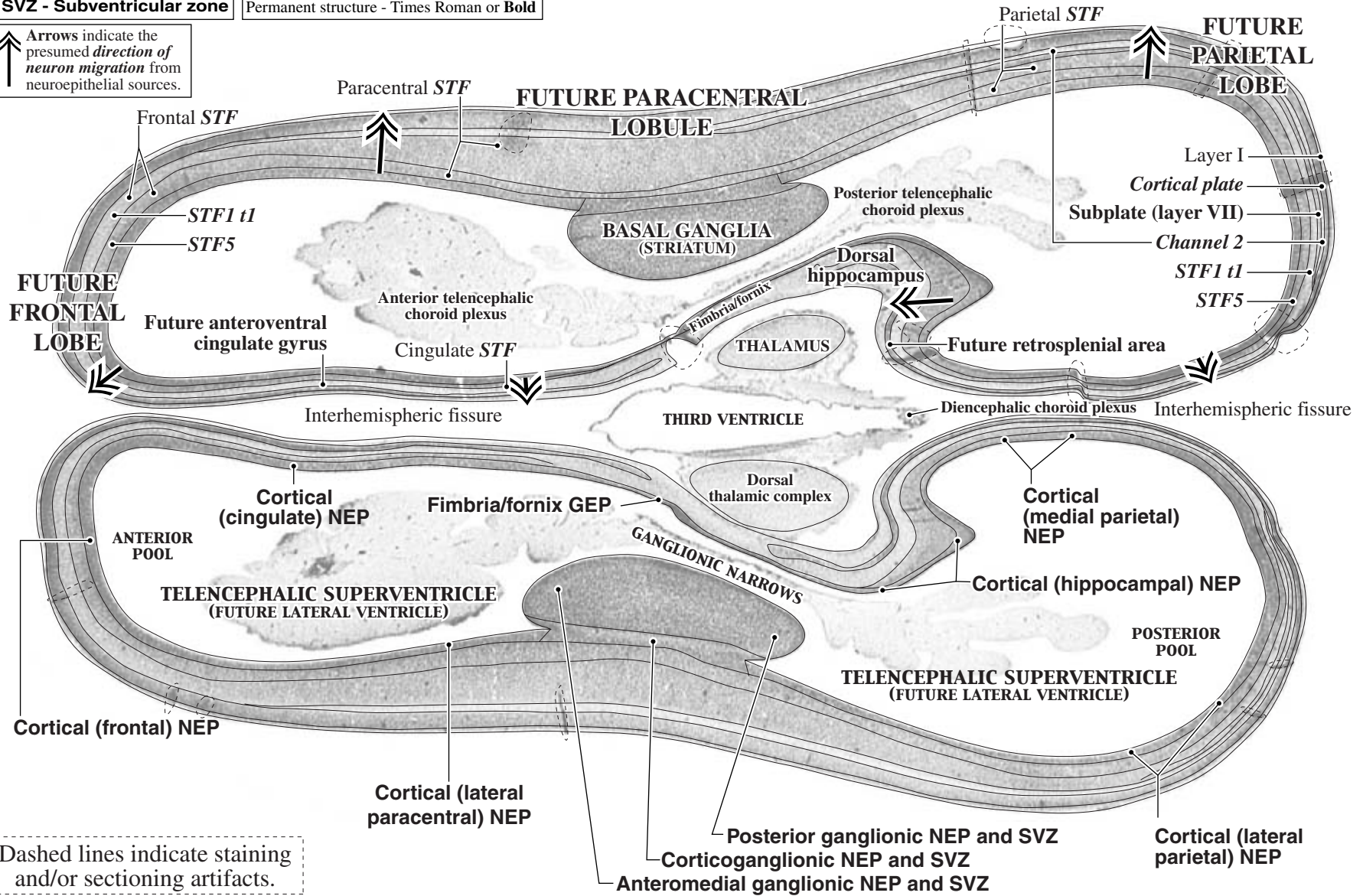
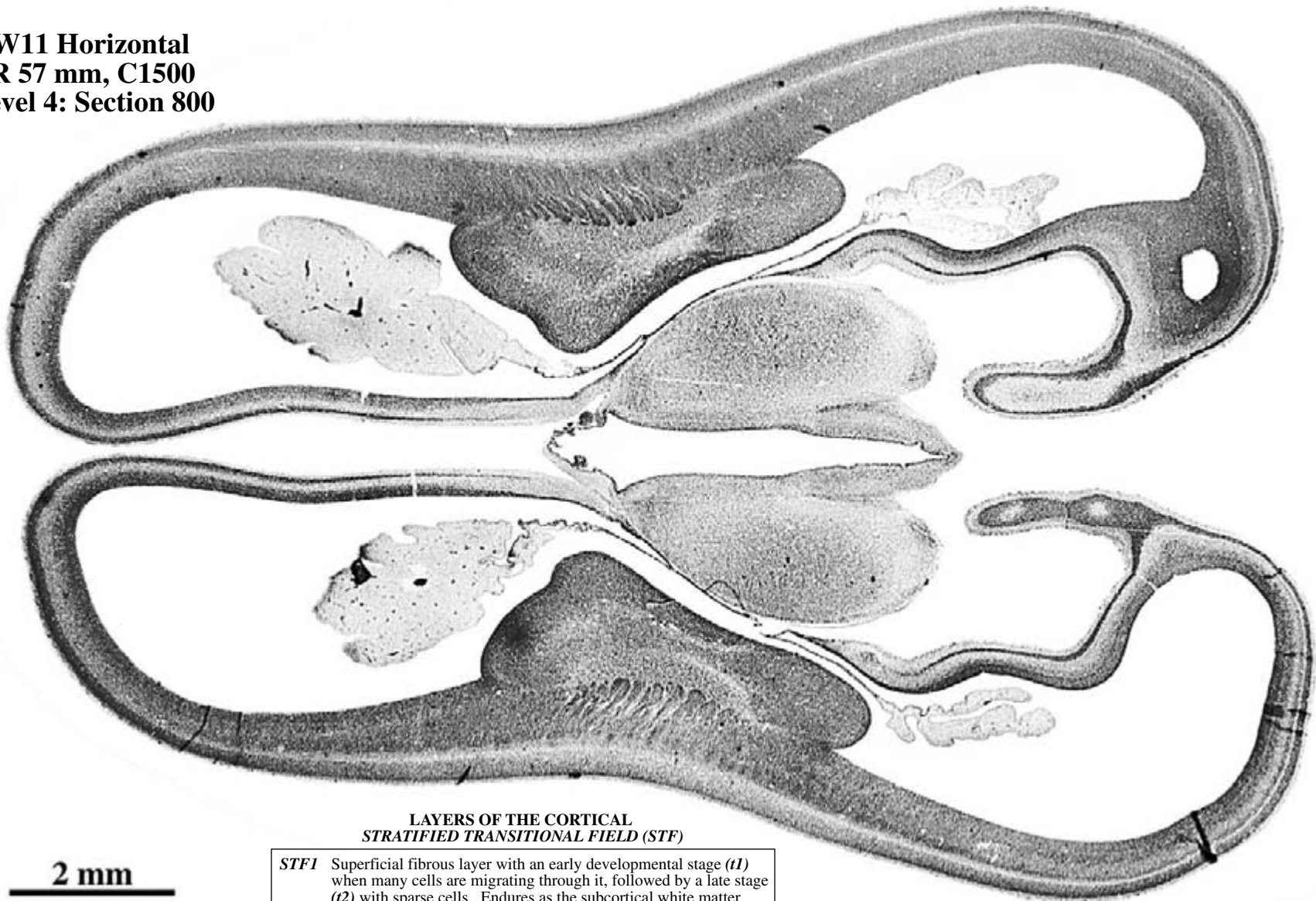


PLATE 39A

**GW11 Horizontal
CR 57 mm, C1500
Level 4: Section 800**



2 mm

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*I1*) when many cells are migrating through it, followed by a late stage (*I2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 39B

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.

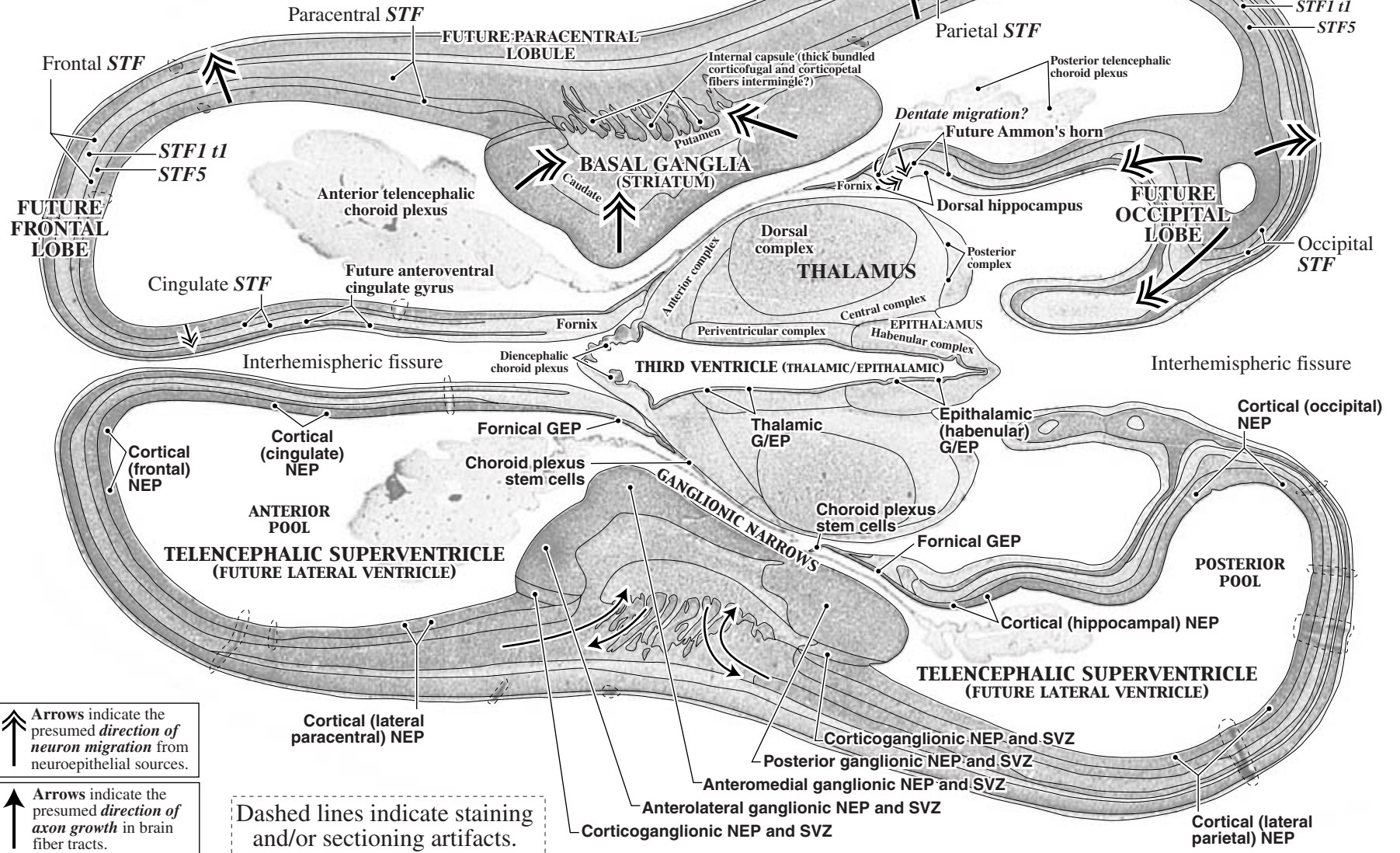
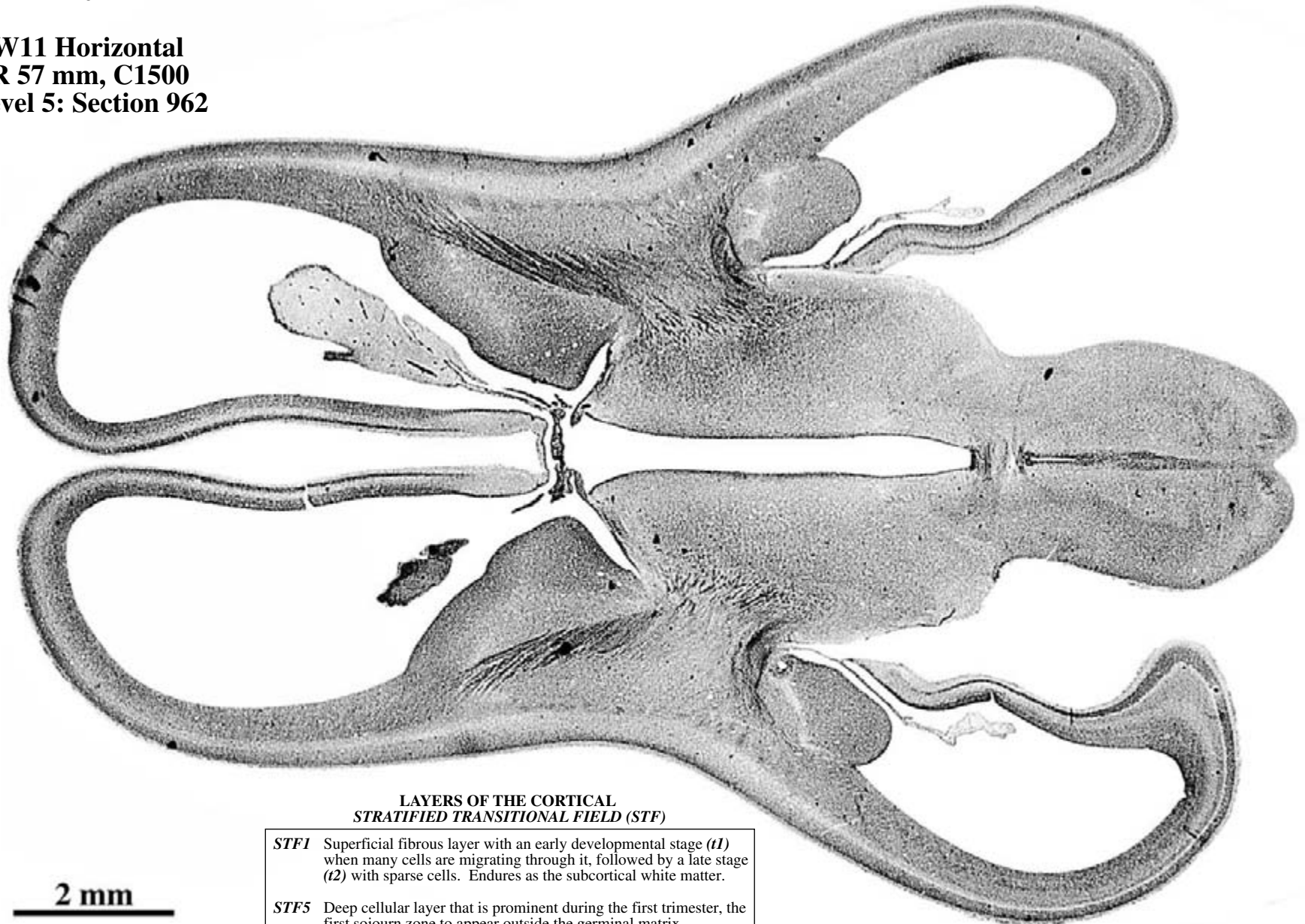


PLATE 40A

**GW11 Horizontal
CR 57 mm, C1500
Level 5: Section 962**



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

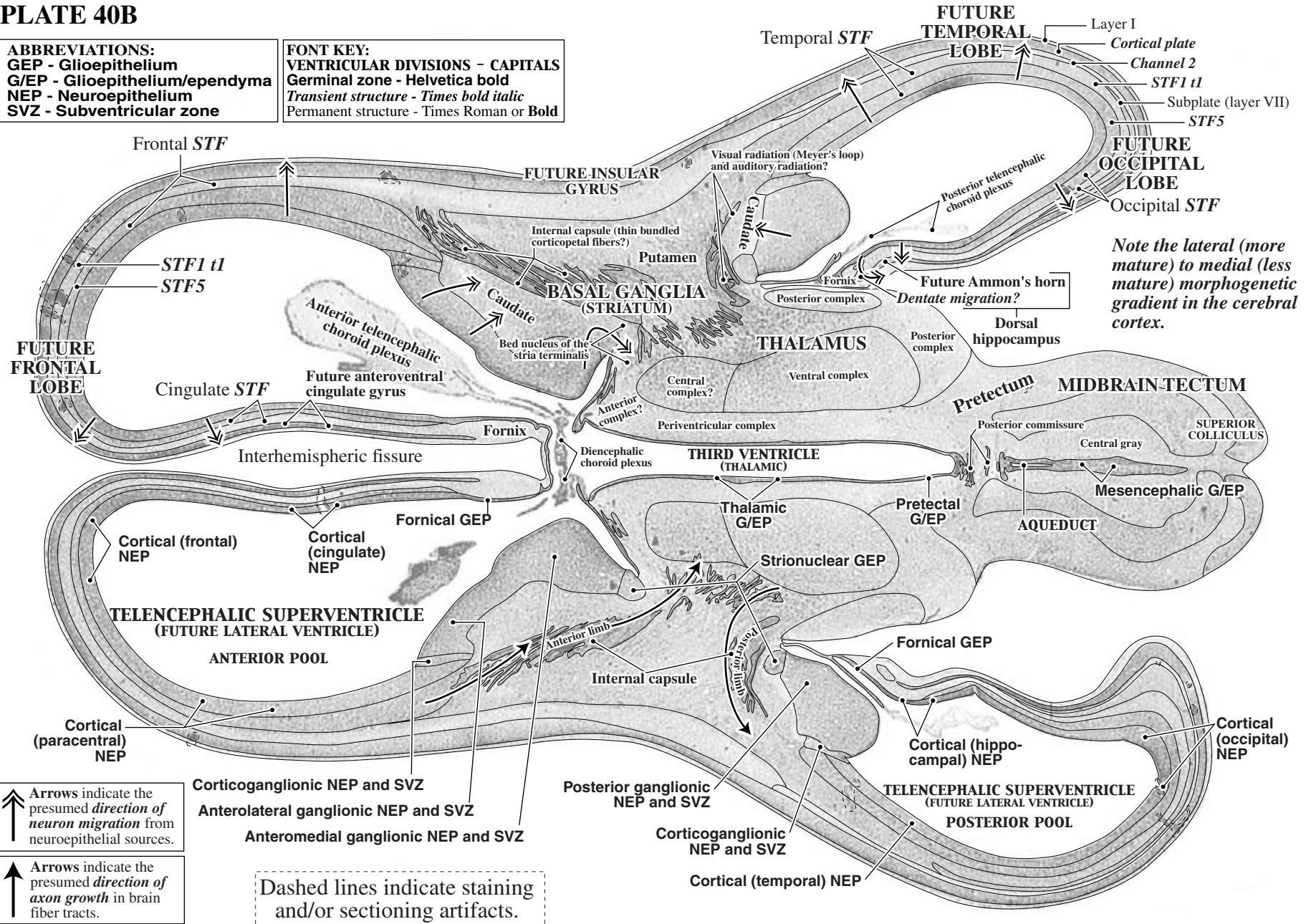
- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

PLATE 40B

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold



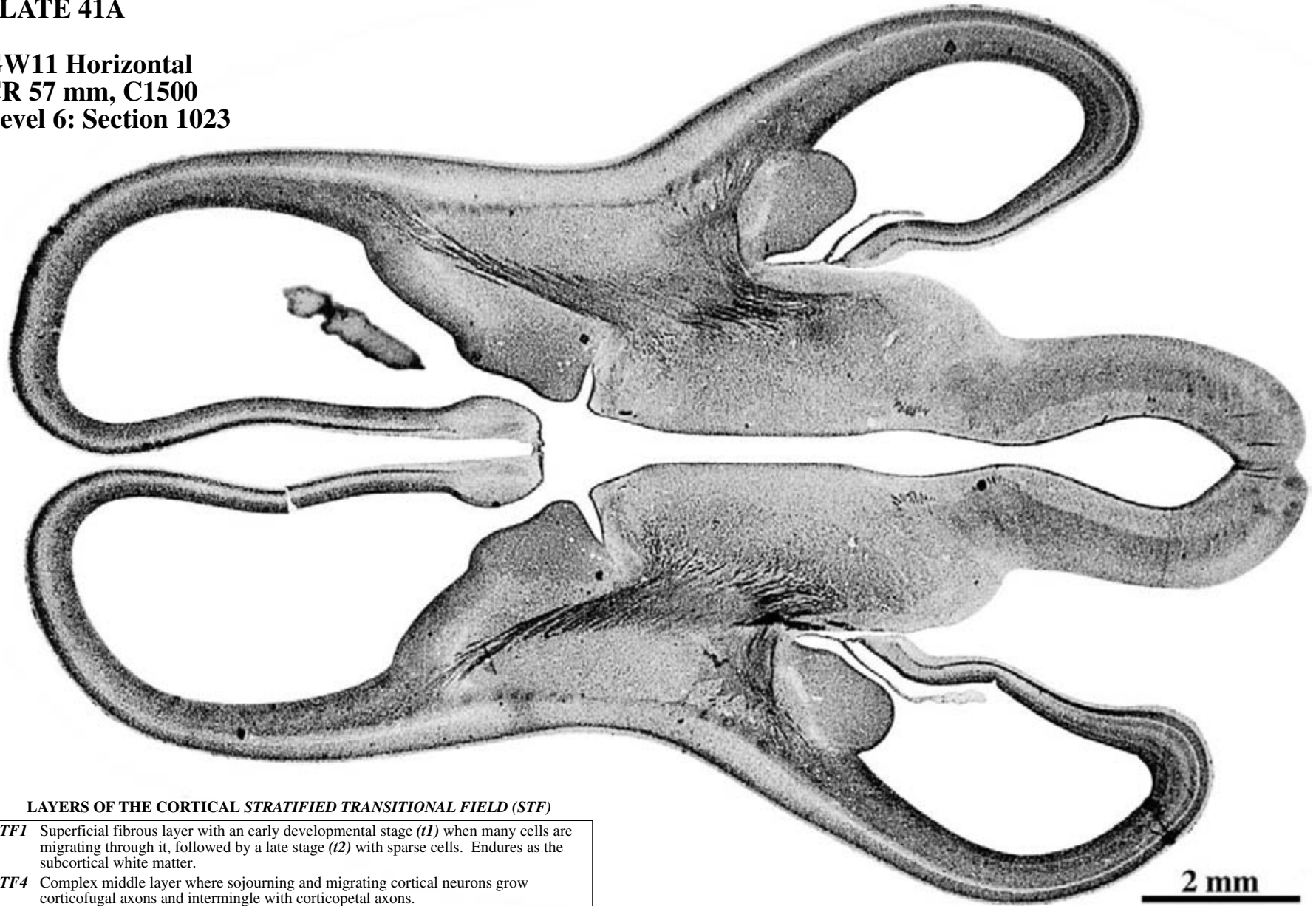
↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

↑ Arrows indicate the presumed direction of axon growth in brain fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 41A

**GW11 Horizontal
CR 57 mm, C1500
Level 6: Section 1023**



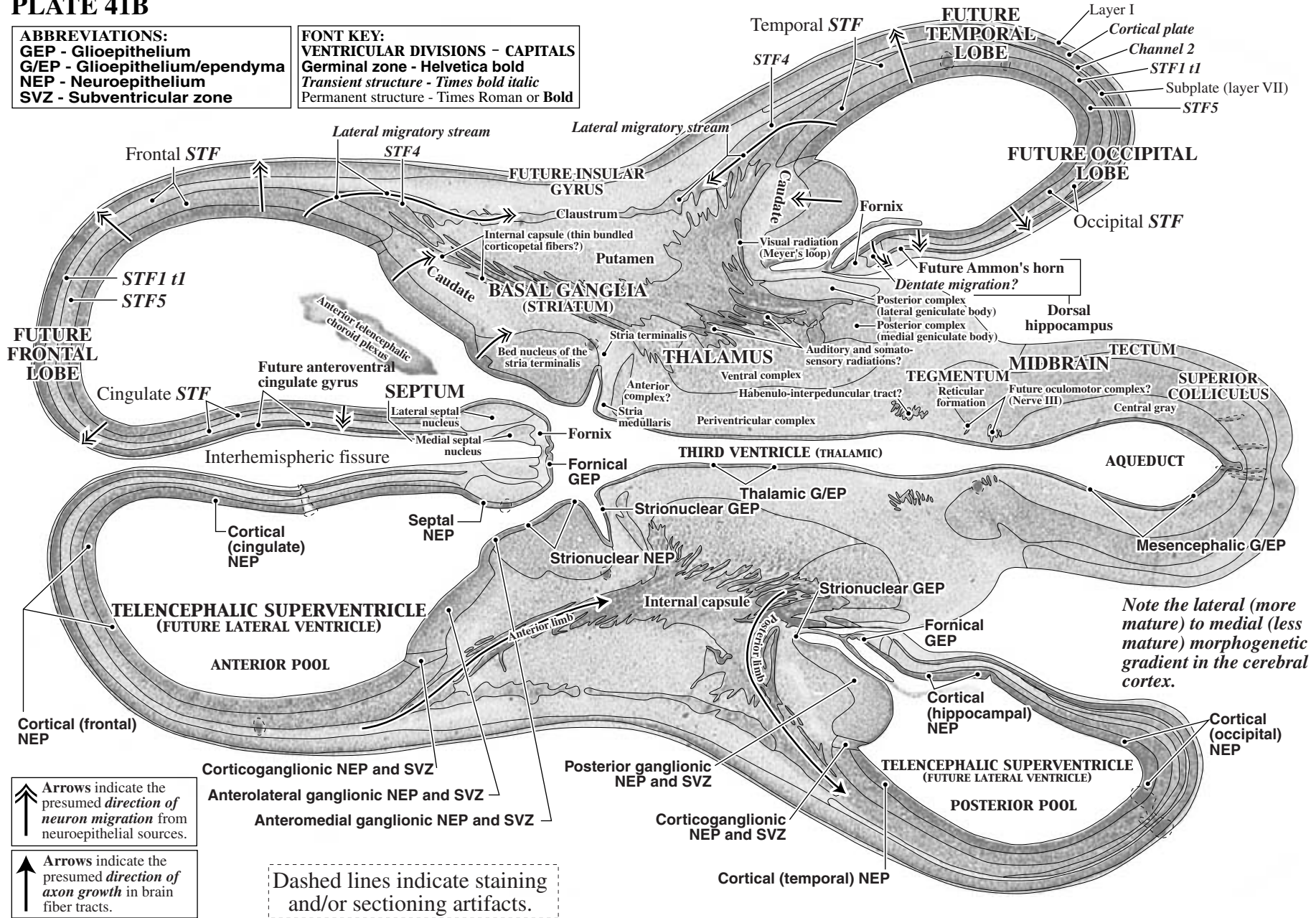
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*I1*) when many cells are migrating through it, followed by a late stage (*I2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 41B

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold



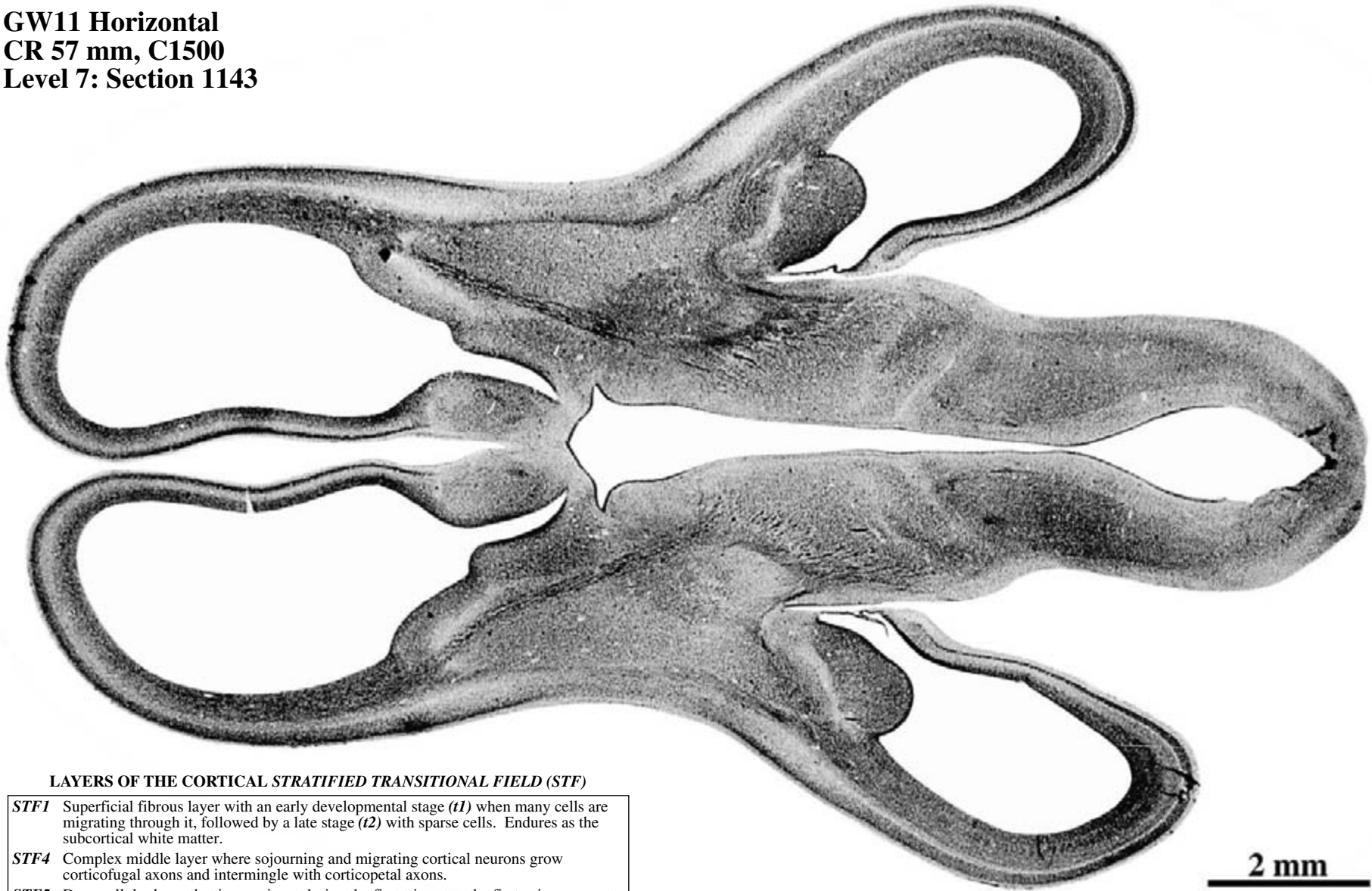
↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

↑ Arrows indicate the presumed direction of axon growth in brain fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 42A

GW11 Horizontal
CR 57 mm, C1500
Level 7: Section 1143



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

PLATE 42B

ABBREVIATIONS:
GEP - Glioeptihelium
G/EP - Glioeptihelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS – CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times Roman or Italic
Permanent structure - Times Roman or Bold

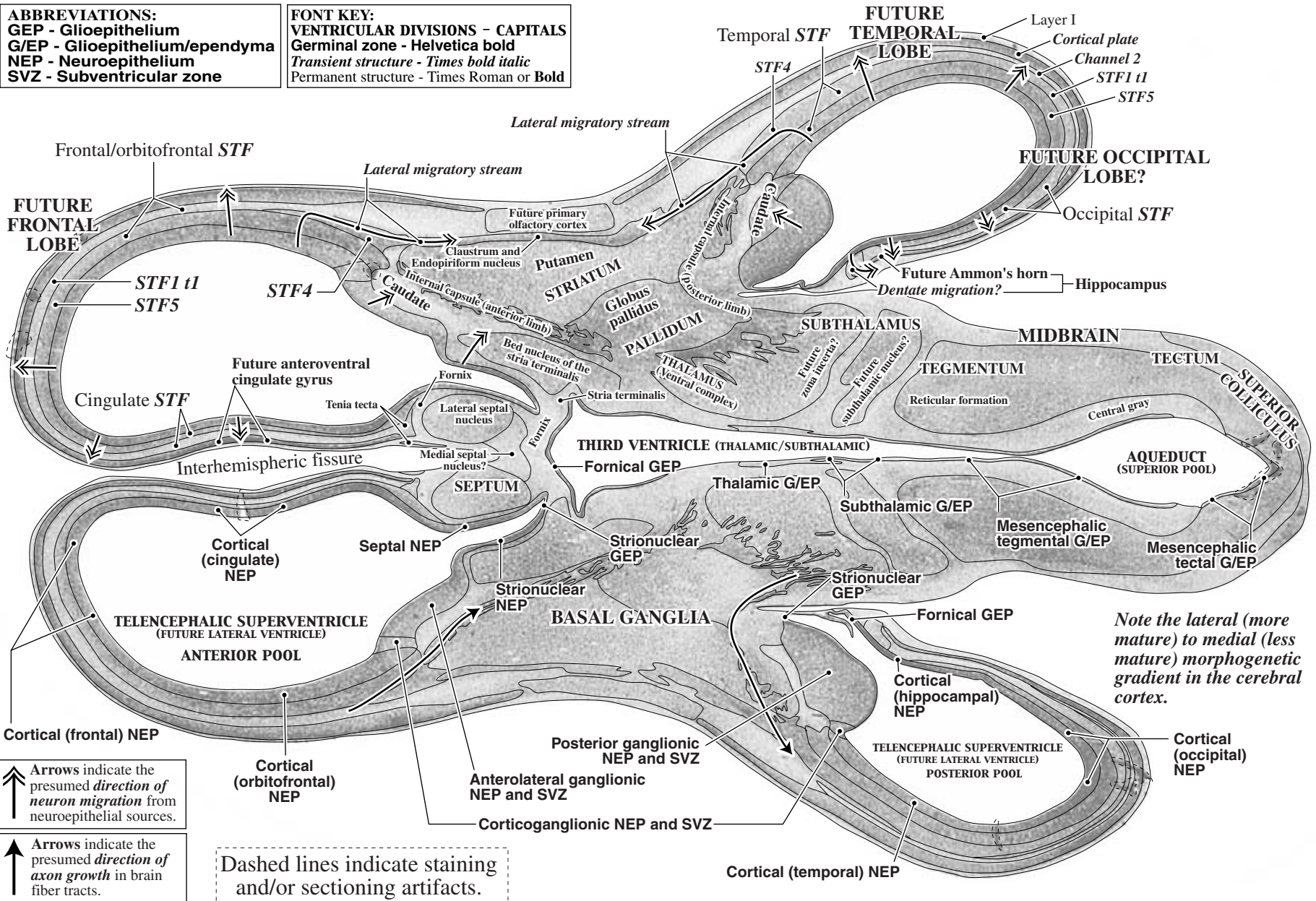
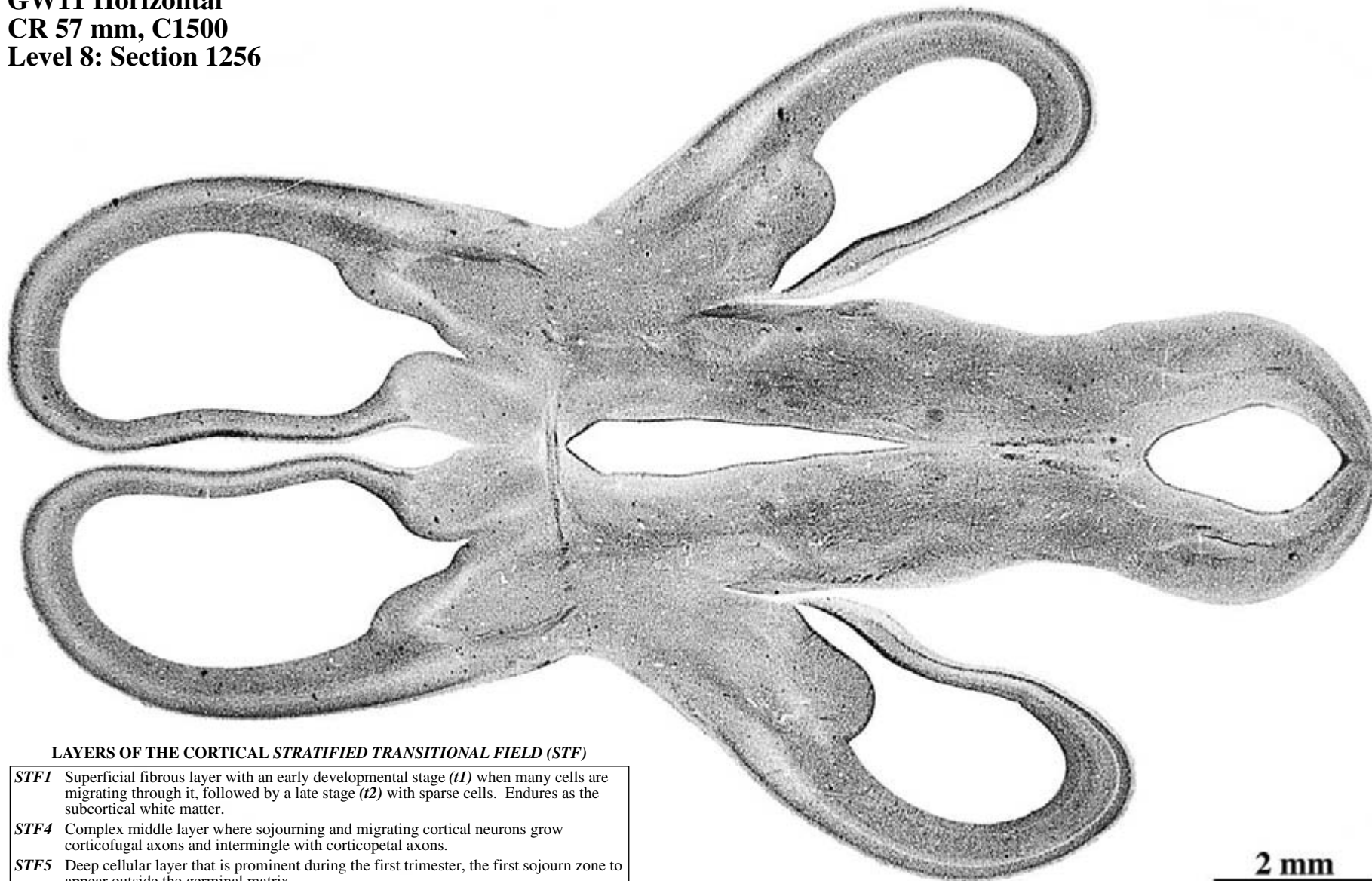


PLATE 43A

GW11 Horizontal
CR 57 mm, C1500
Level 8: Section 1256



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

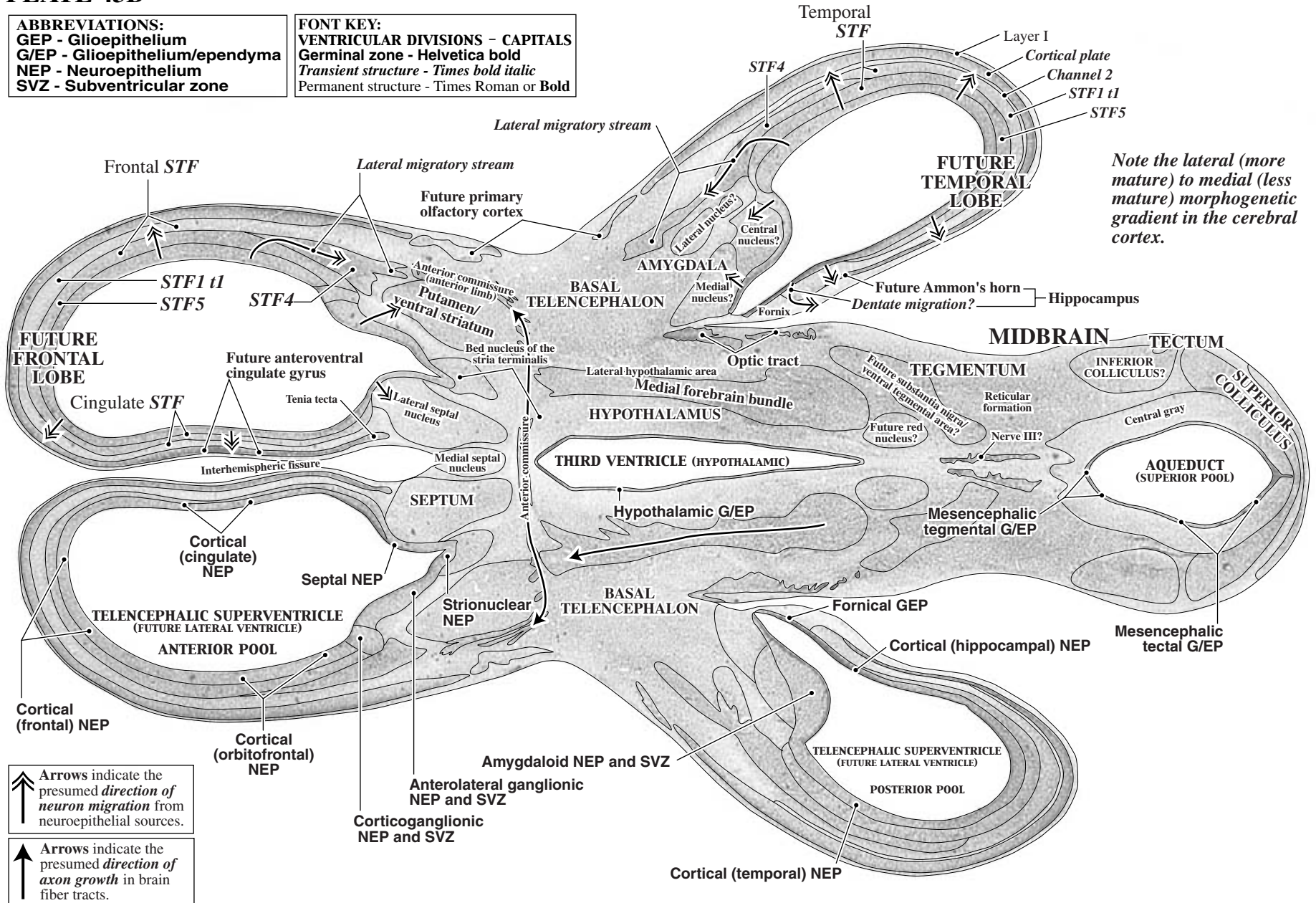
- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

PLATE 43B

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



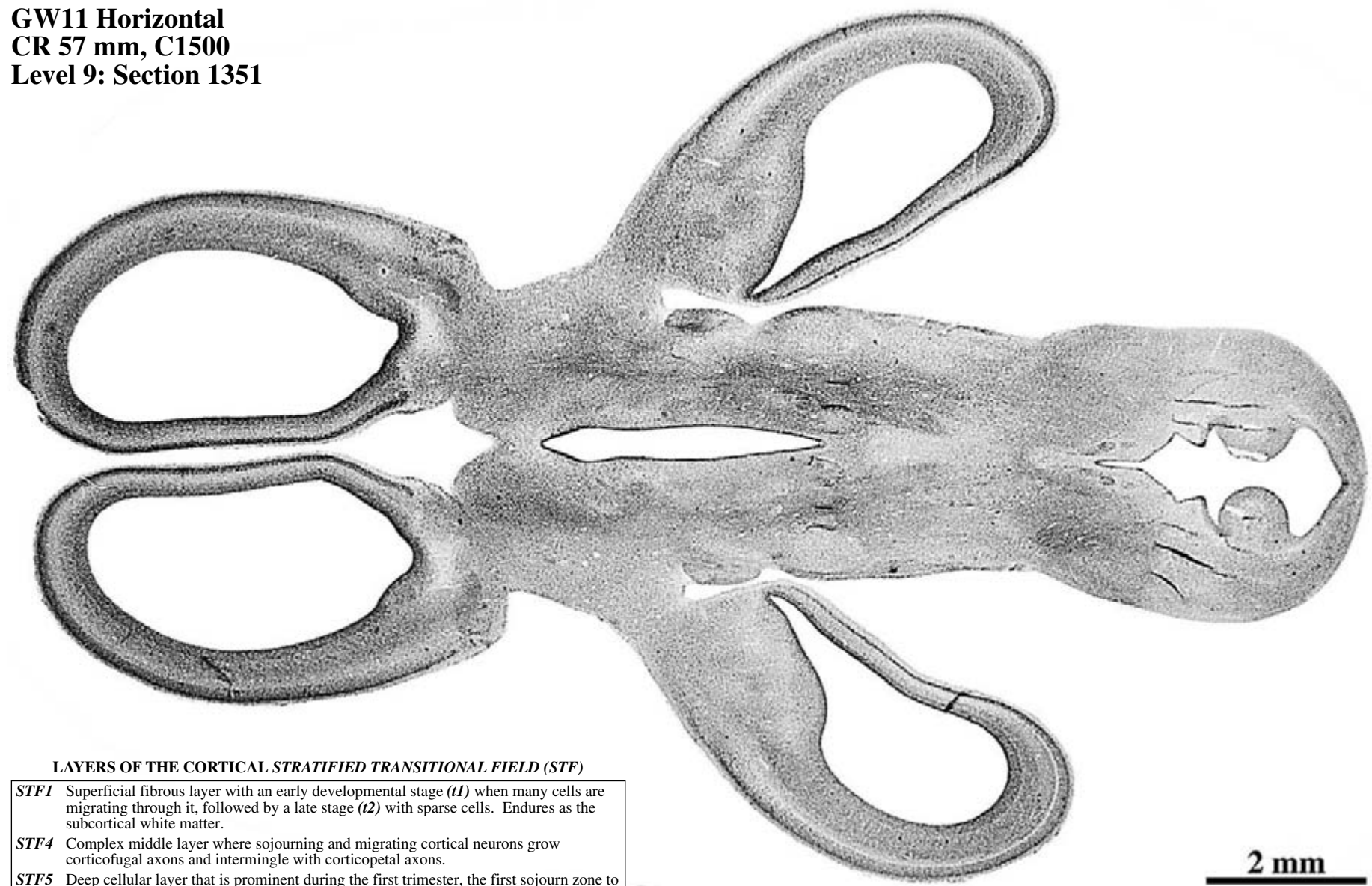
Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 44A

**GW11 Horizontal
CR 57 mm, C1500
Level 9: Section 1351**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

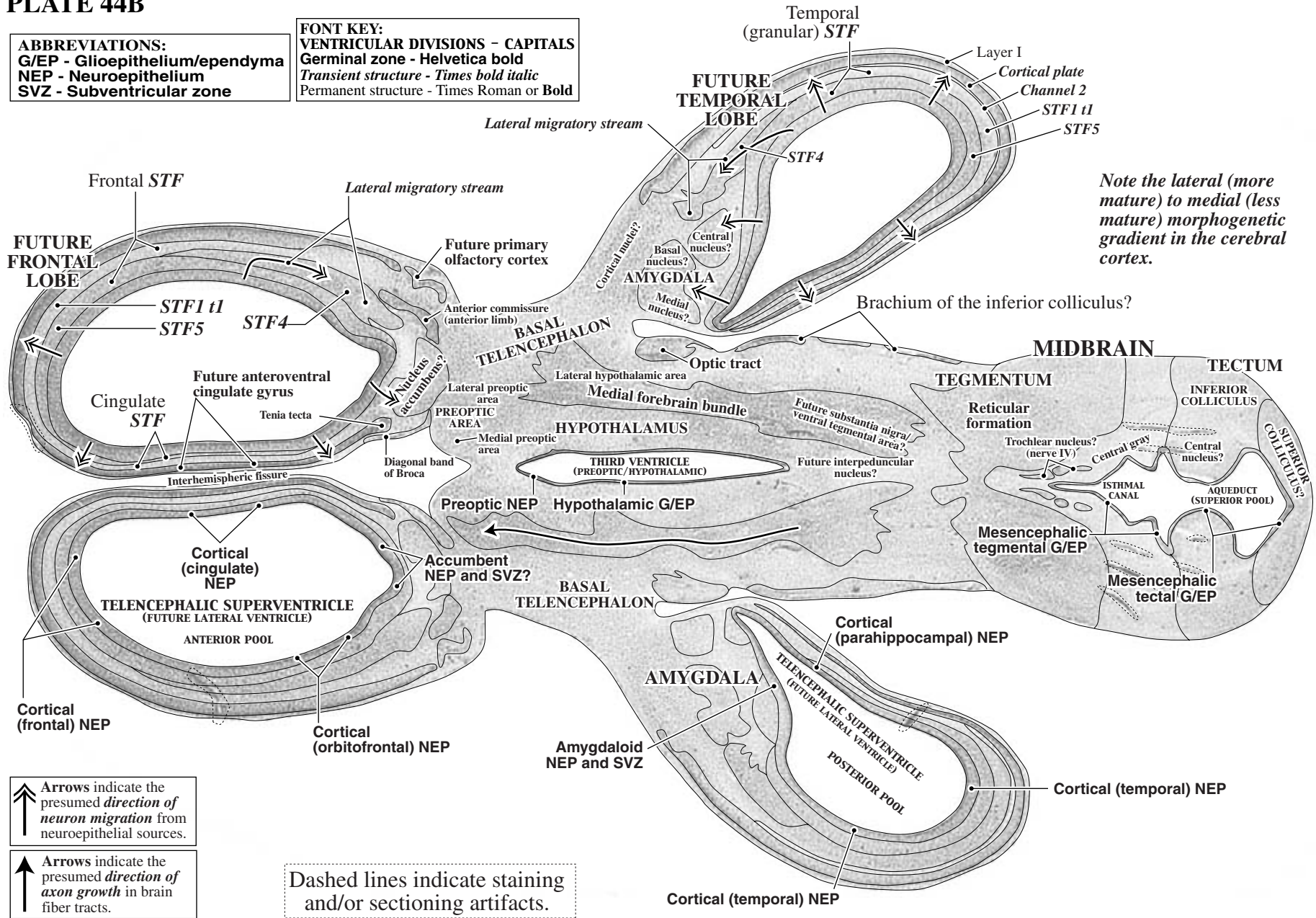
- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

PLATE 44B

ABBREVIATIONS:
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.

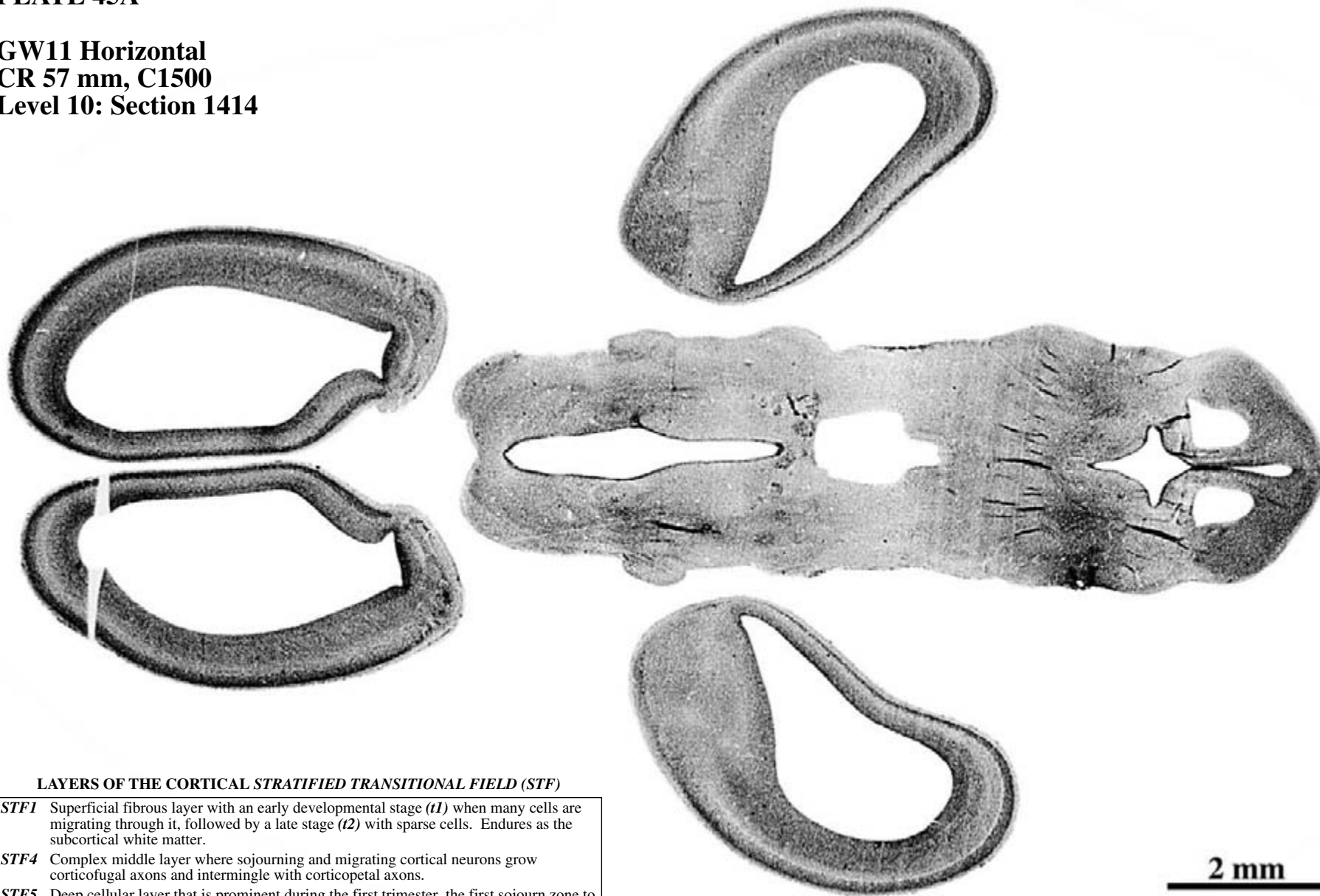
↑ Arrows indicate the presumed **direction of neuron migration** from neuroepithelial sources.

↑ Arrows indicate the presumed **direction of axon growth** in brain fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 45A

GW11 Horizontal
CR 57 mm, C1500
Level 10: Section 1414



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

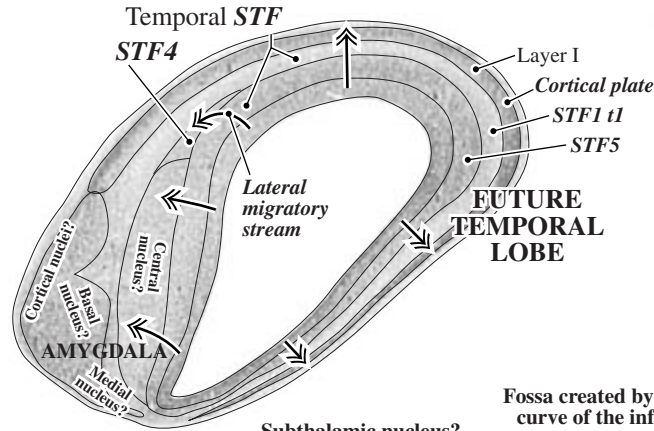
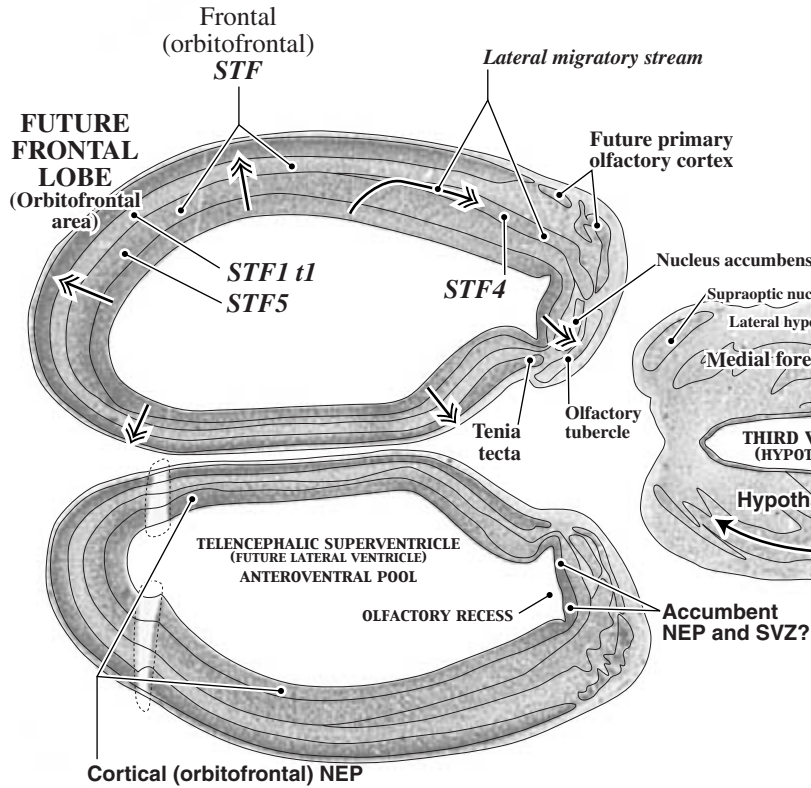
- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

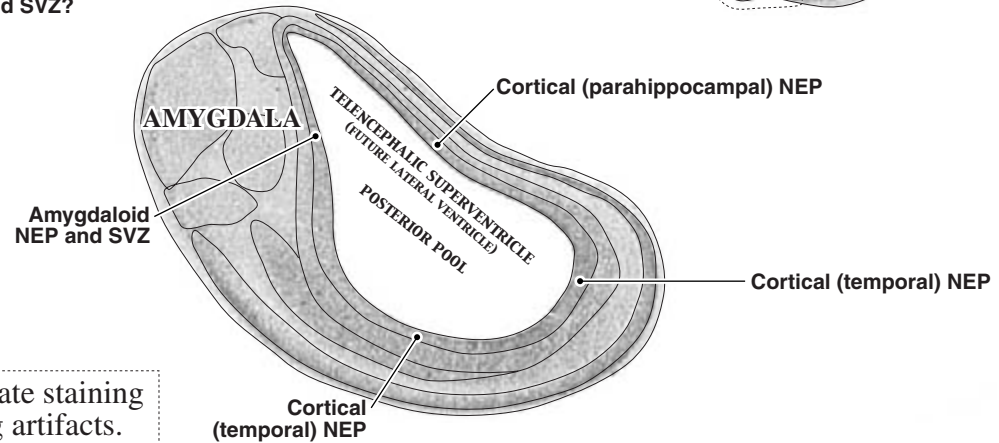
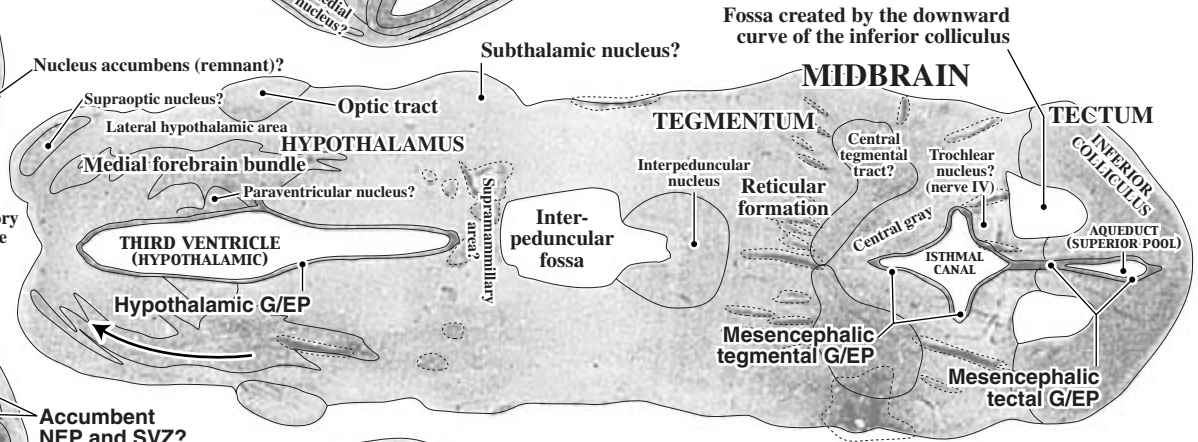
PLATE 45B

ABBREVIATIONS:
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold



Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.



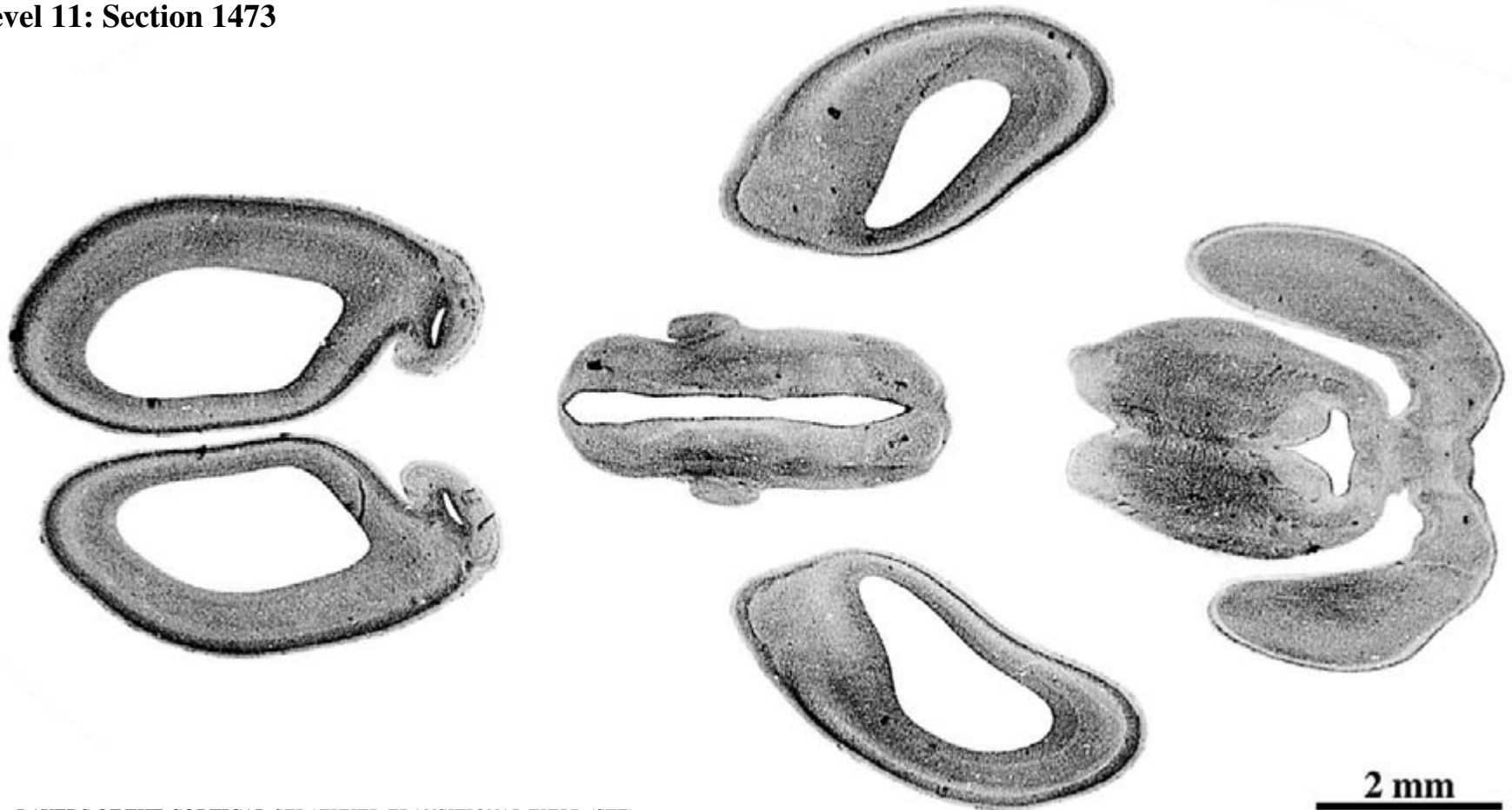
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 46A

**GW11 Horizontal
CR 57 mm, C1500
Level 11: Section 1473**



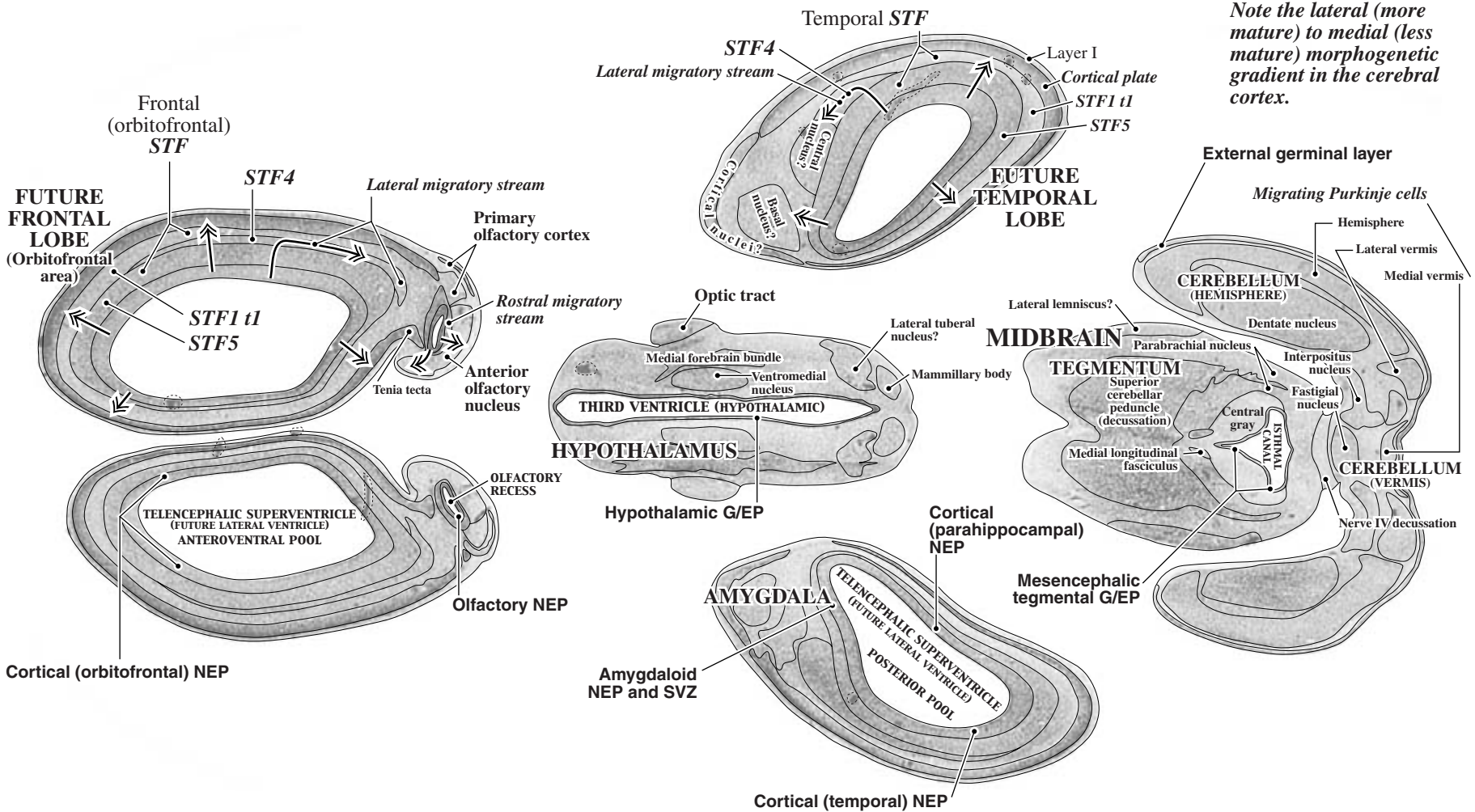
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 46B

ABBREVIATIONS:
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



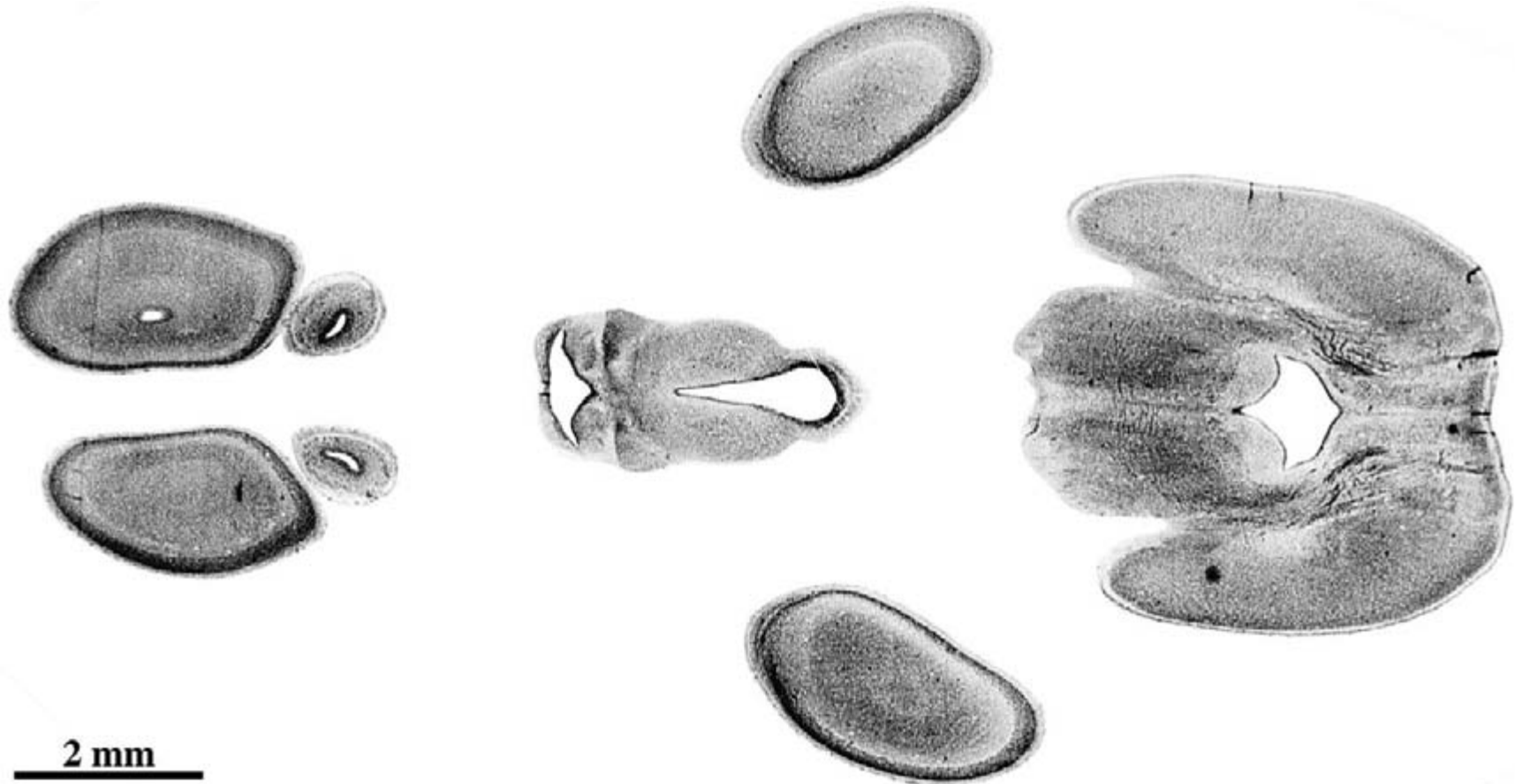
Note the lateral (more mature) to medial (less mature) morphogenetic gradient in the cerebral cortex.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 47A

**GW11 Horizontal
CR 57 mm, C1500
Level 12: Section 1566**



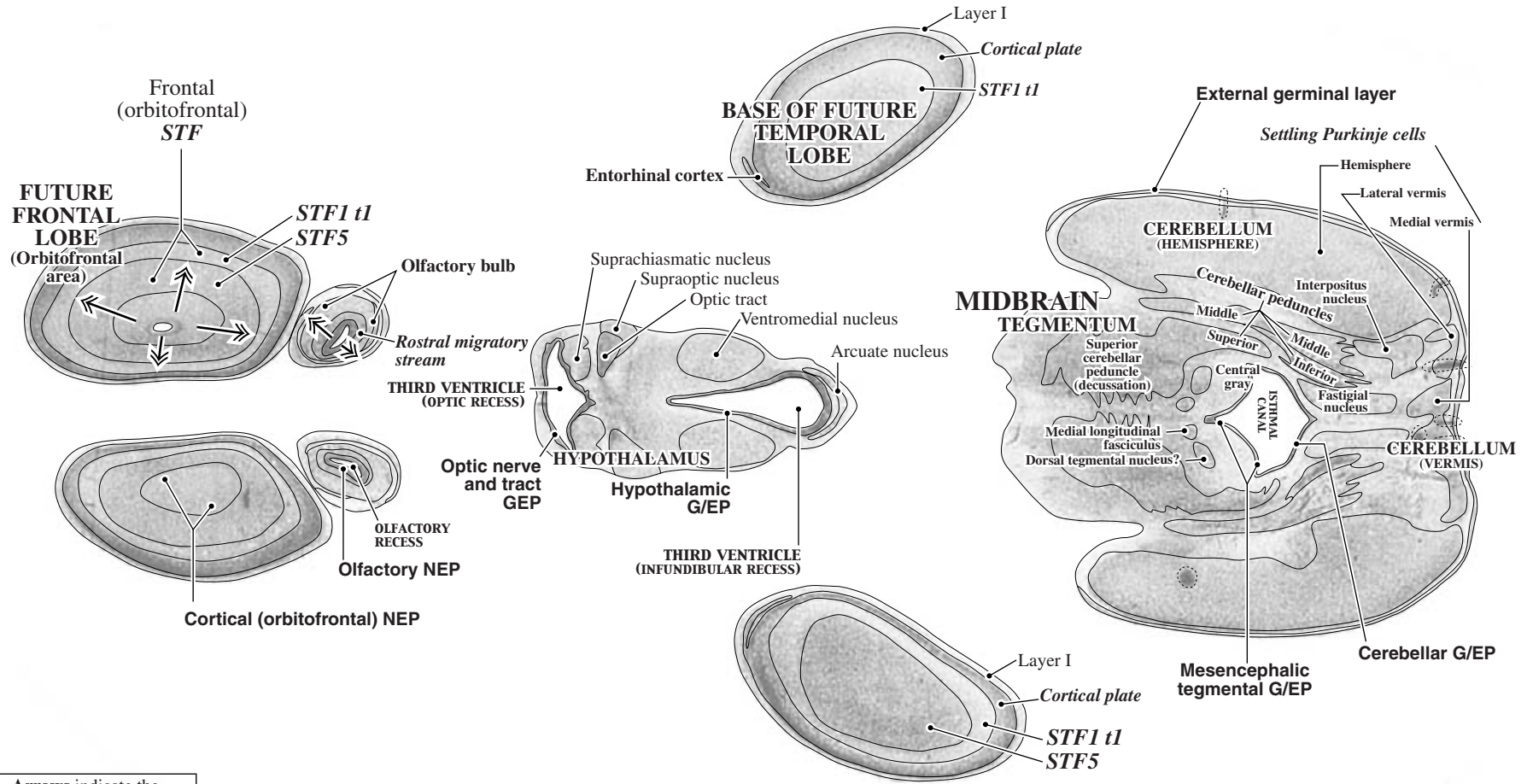
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 47B

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 48A

**GW11 Horizontal
CR 57 mm, C1500
Level 13: Section 1626**

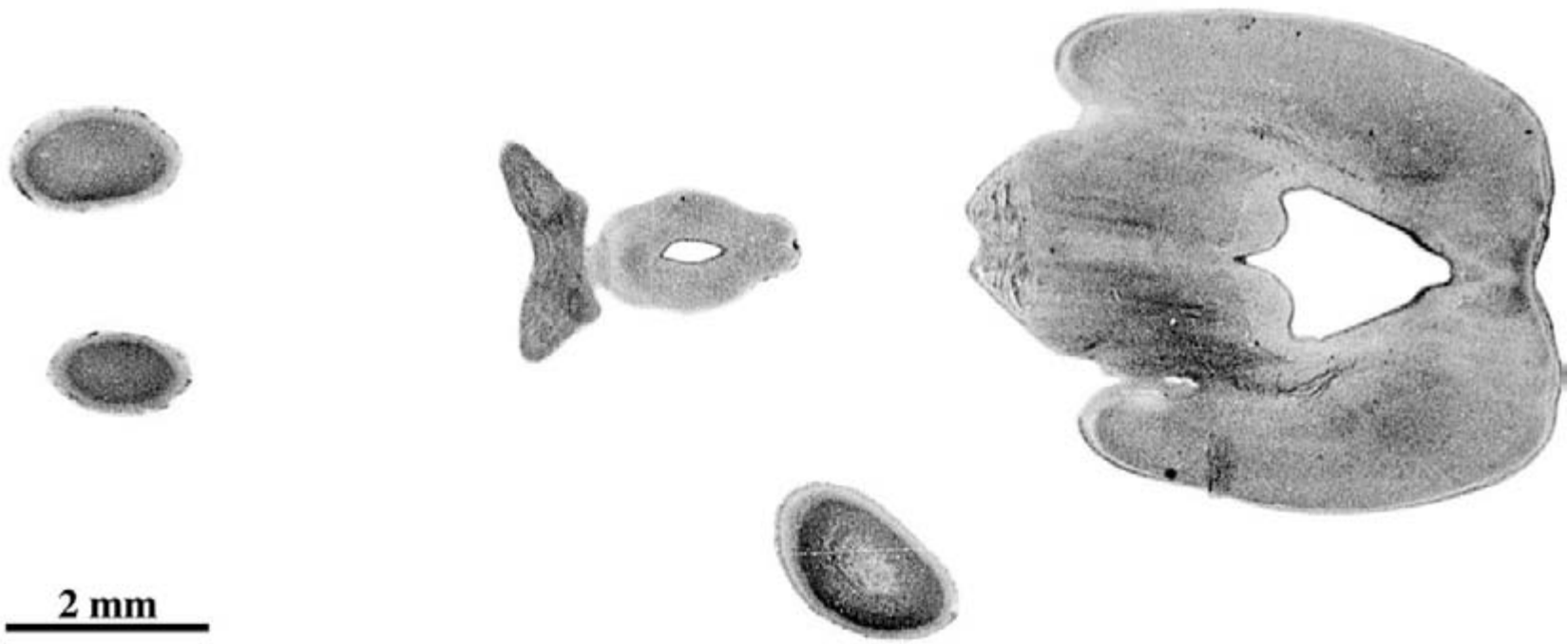
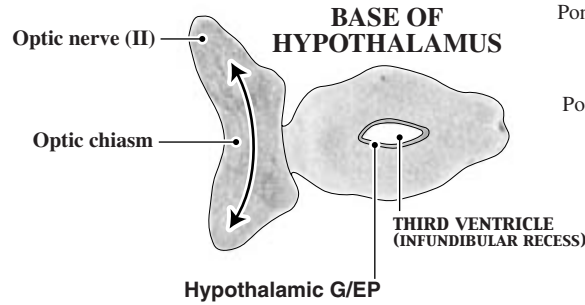
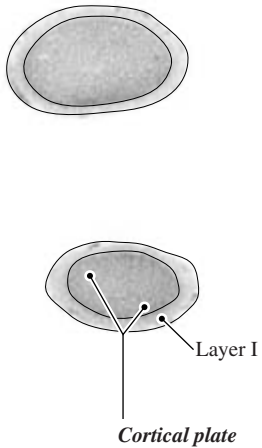


PLATE 48B

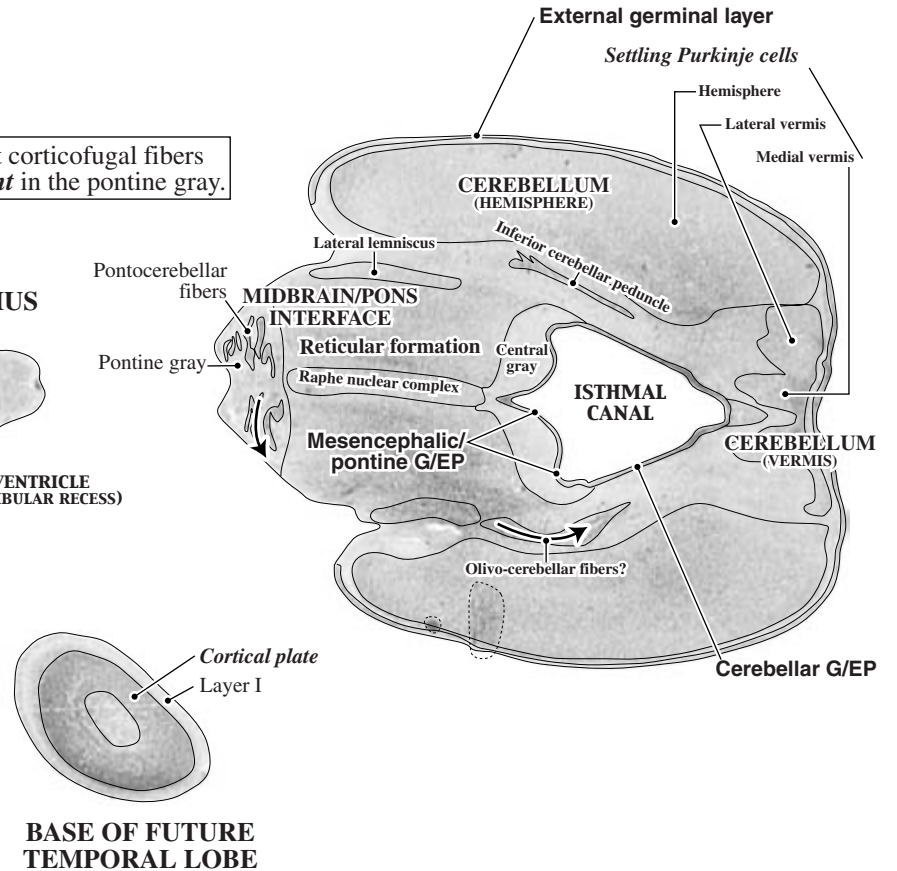
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioeptihelium/ependyma

**BASE OF FUTURE
FRONTAL LOBE
(Orbitofrontal area)**



Note that corticofugal fibers are *absent* in the pontine gray.

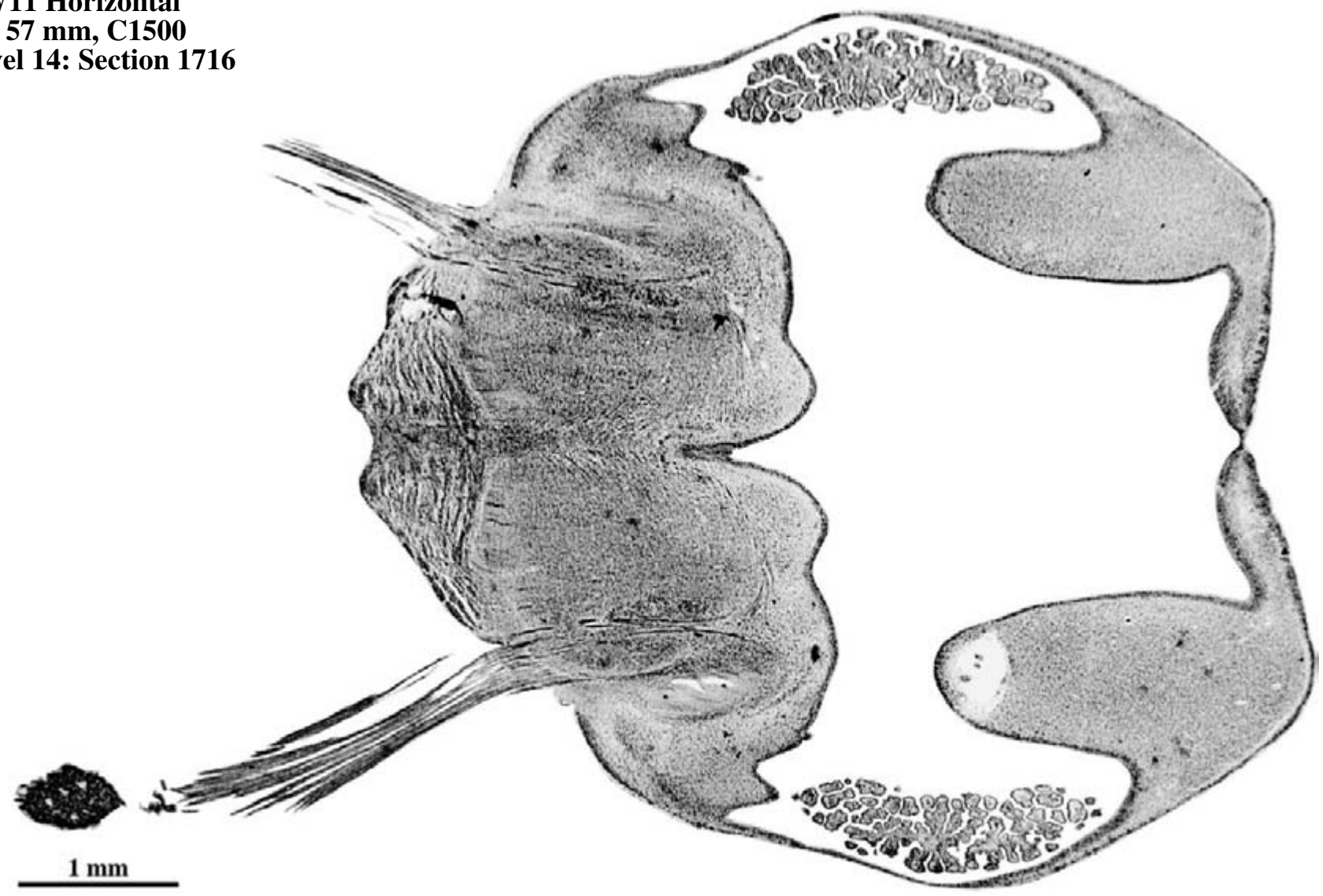


↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 49A

**GW11 Horizontal
CR 57 mm, C1500
Level 14: Section 1716**



Levels 14 to 21 are only shown at high magnification.

PLATE 49B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium

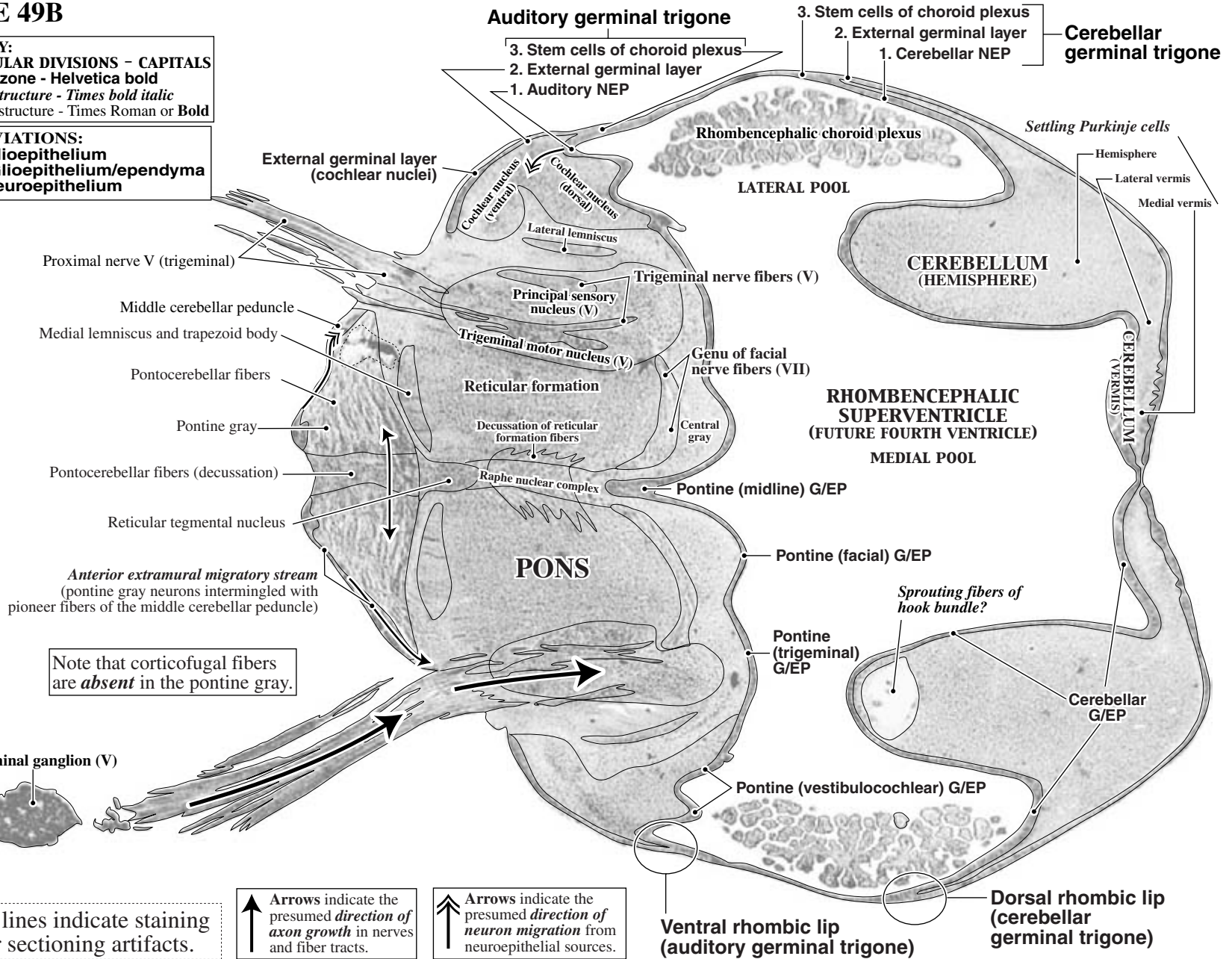
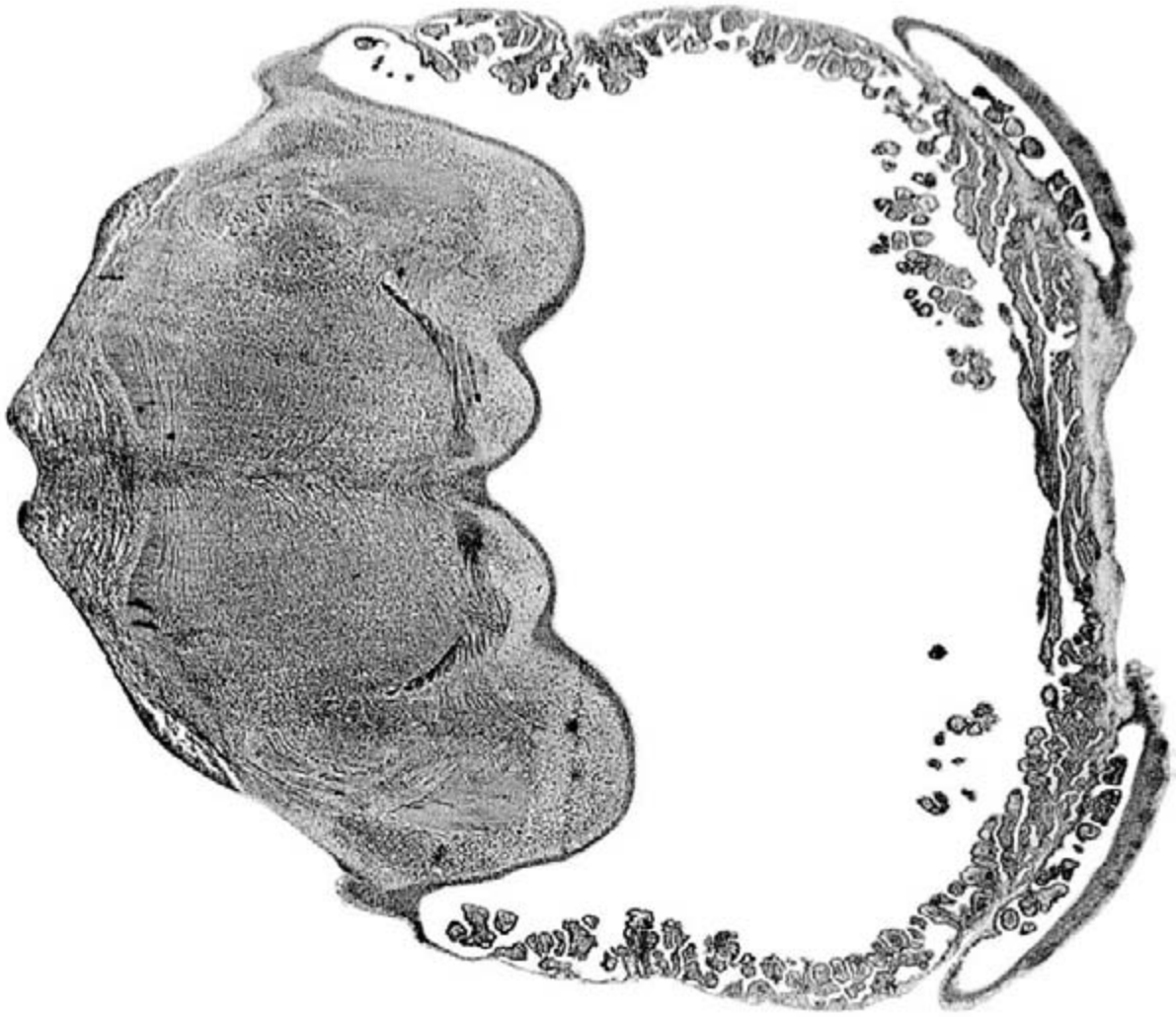


PLATE 50A

**GW11 Horizontal
CR 57 mm, C1500
Level 15: Section 1805**



1 mm

Levels 14 to 21 are only shown at high magnification.

PLATE 50B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**
G/EP - Gliopitthelium/ependyma

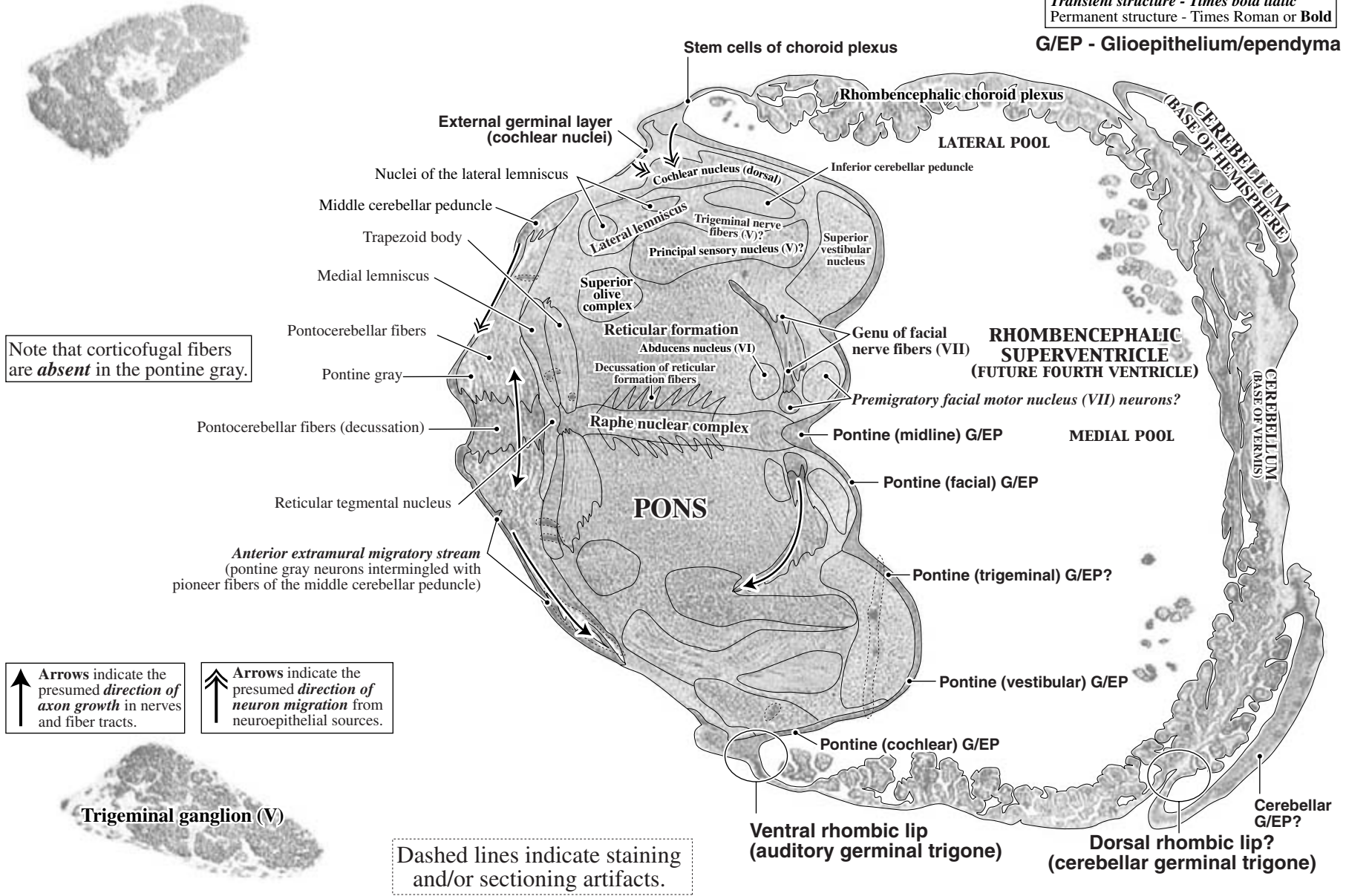
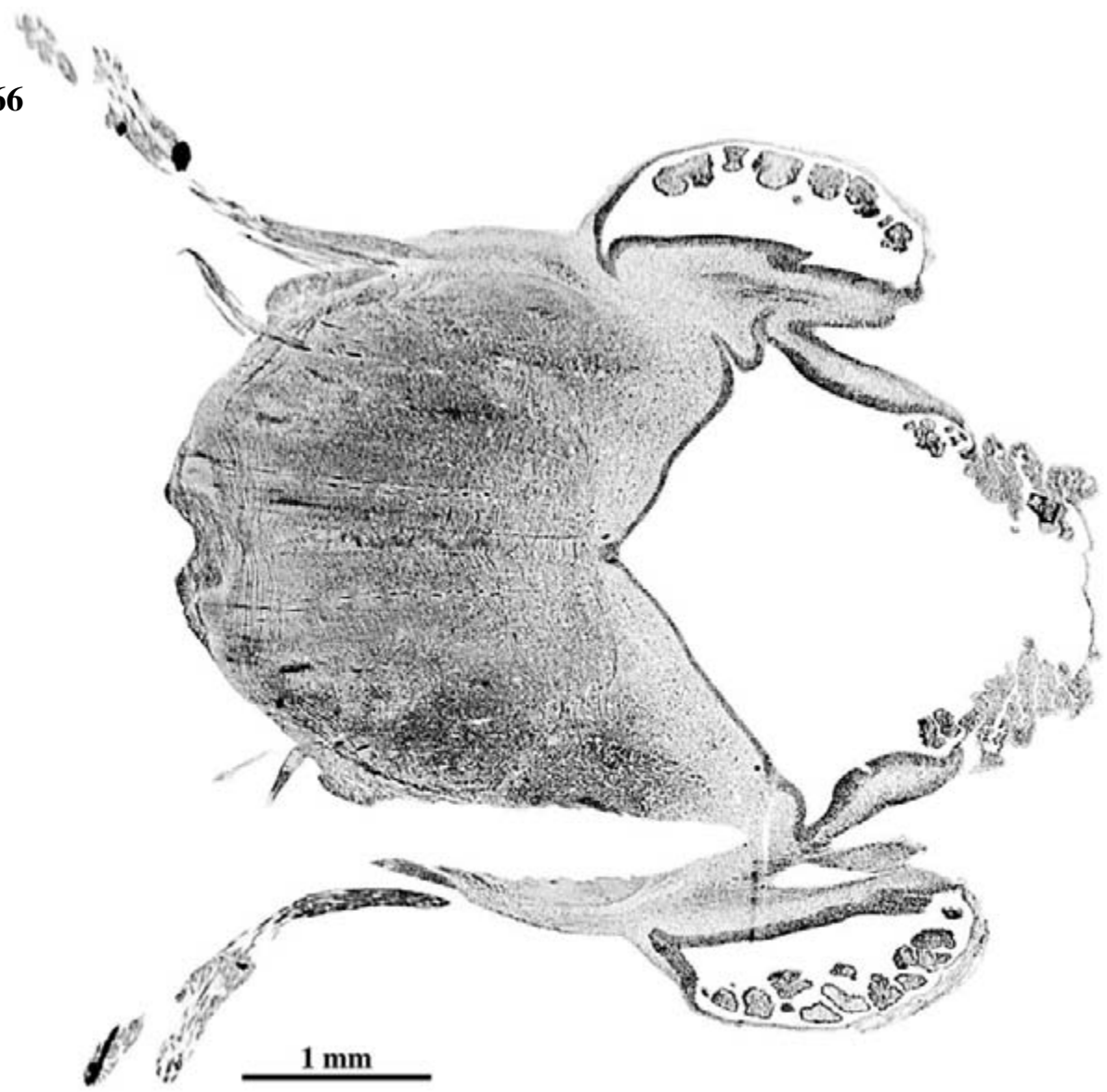


PLATE 51A

**GW11 Horizontal
CR 57 mm, C1500
Level 16: Section 1866**

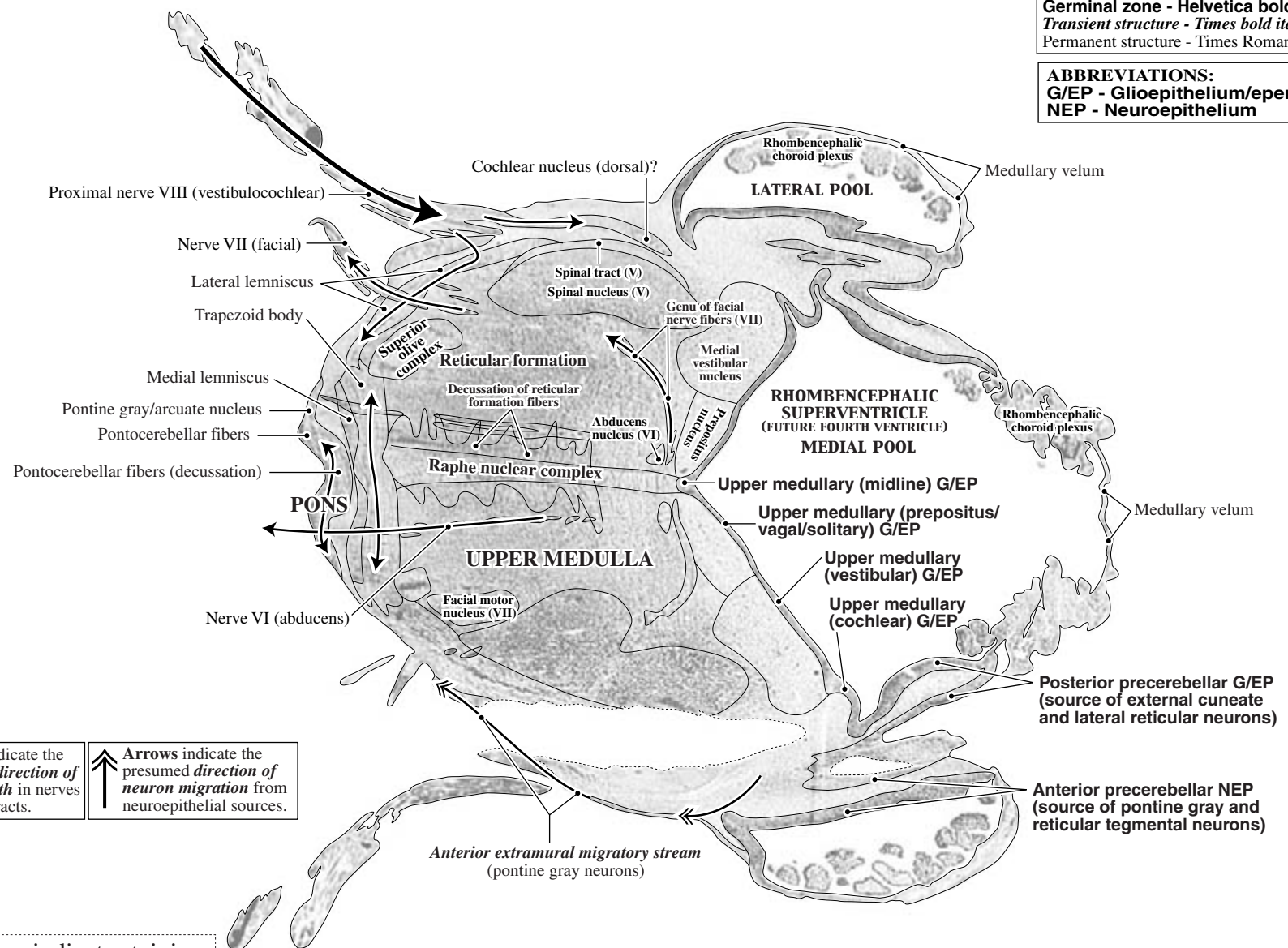


Levels 14 to 21 are only shown at high magnification.

PLATE 51B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium



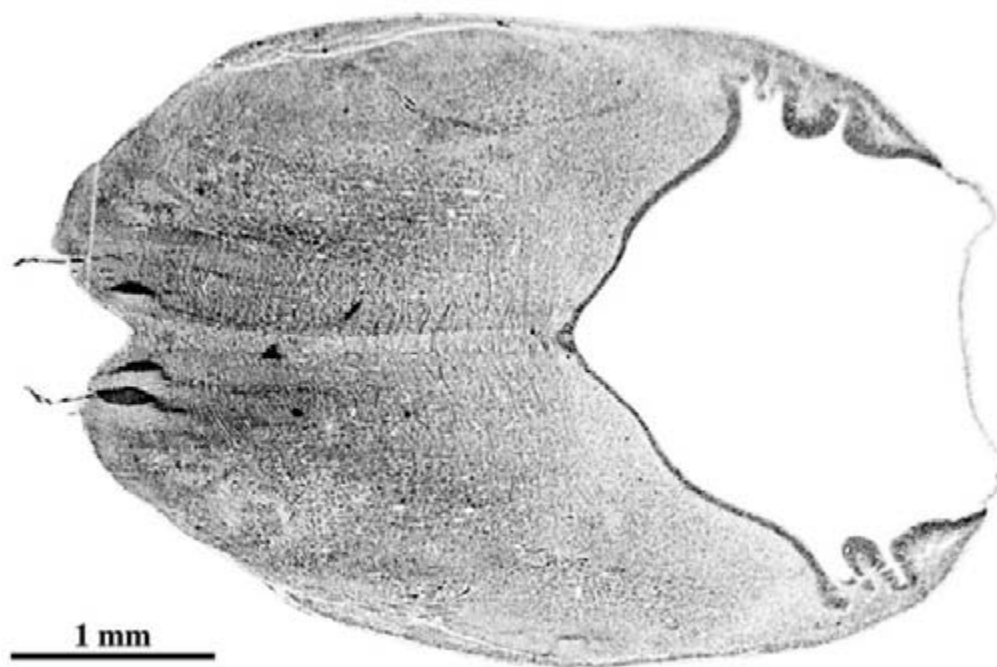
↑ Arrows indicate the presumed *direction of axon growth* in nerves and fiber tracts.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 52A

**GW11 Horizontal
CR 57 mm, C1500
Level 17: Section 1926**

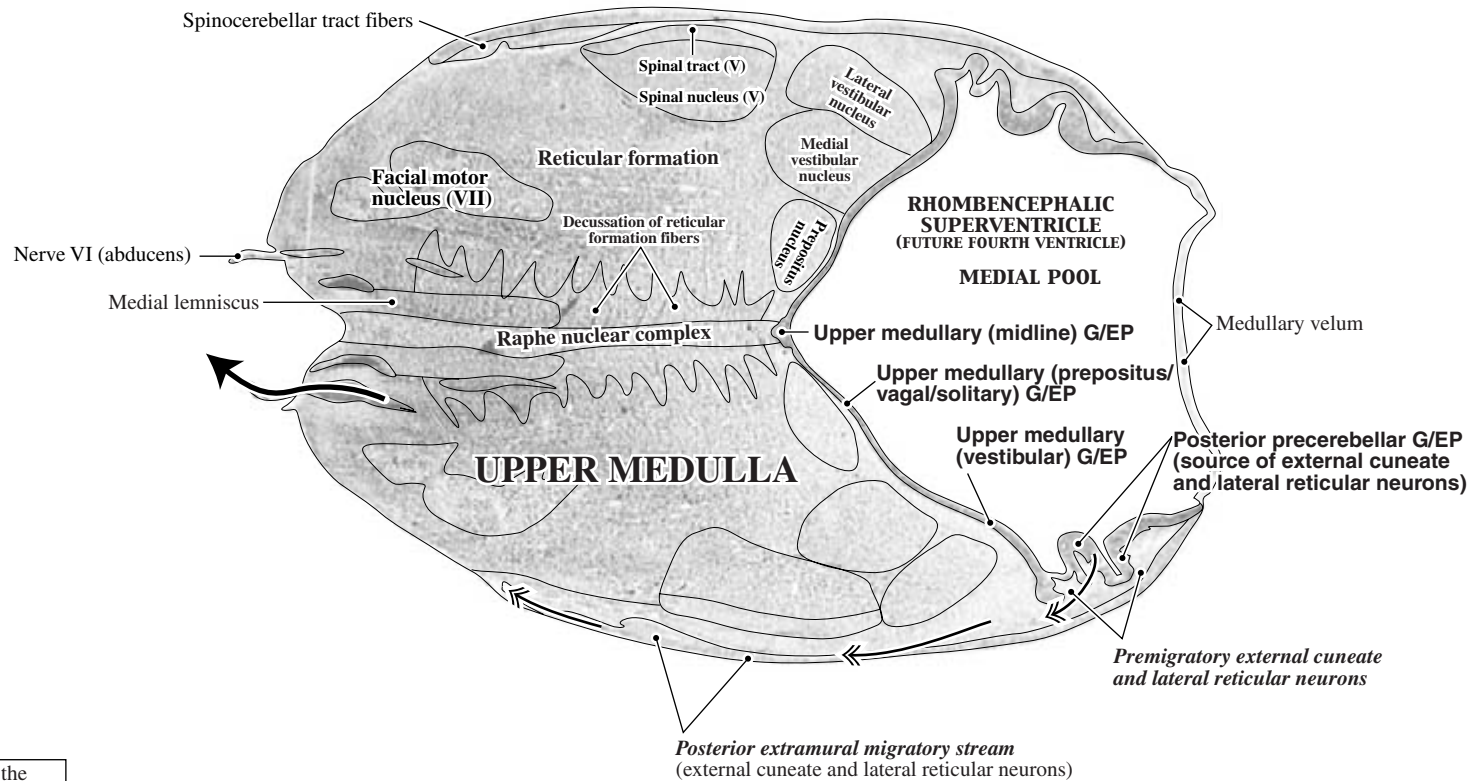


Levels 14 to 21 are only shown at high magnification.

PLATE 52B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioepithelium/ependyma

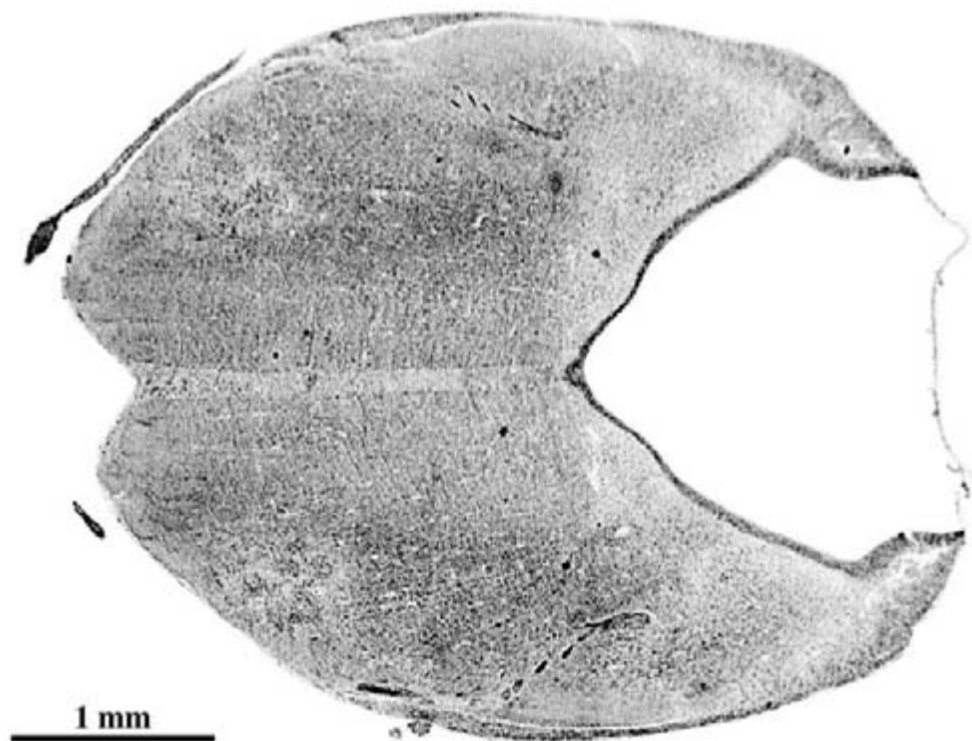


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in nerves and fiber tracts.

PLATE 53A

**GW11 Horizontal
CR 57 mm, C1500
Level 18: Section 1962**

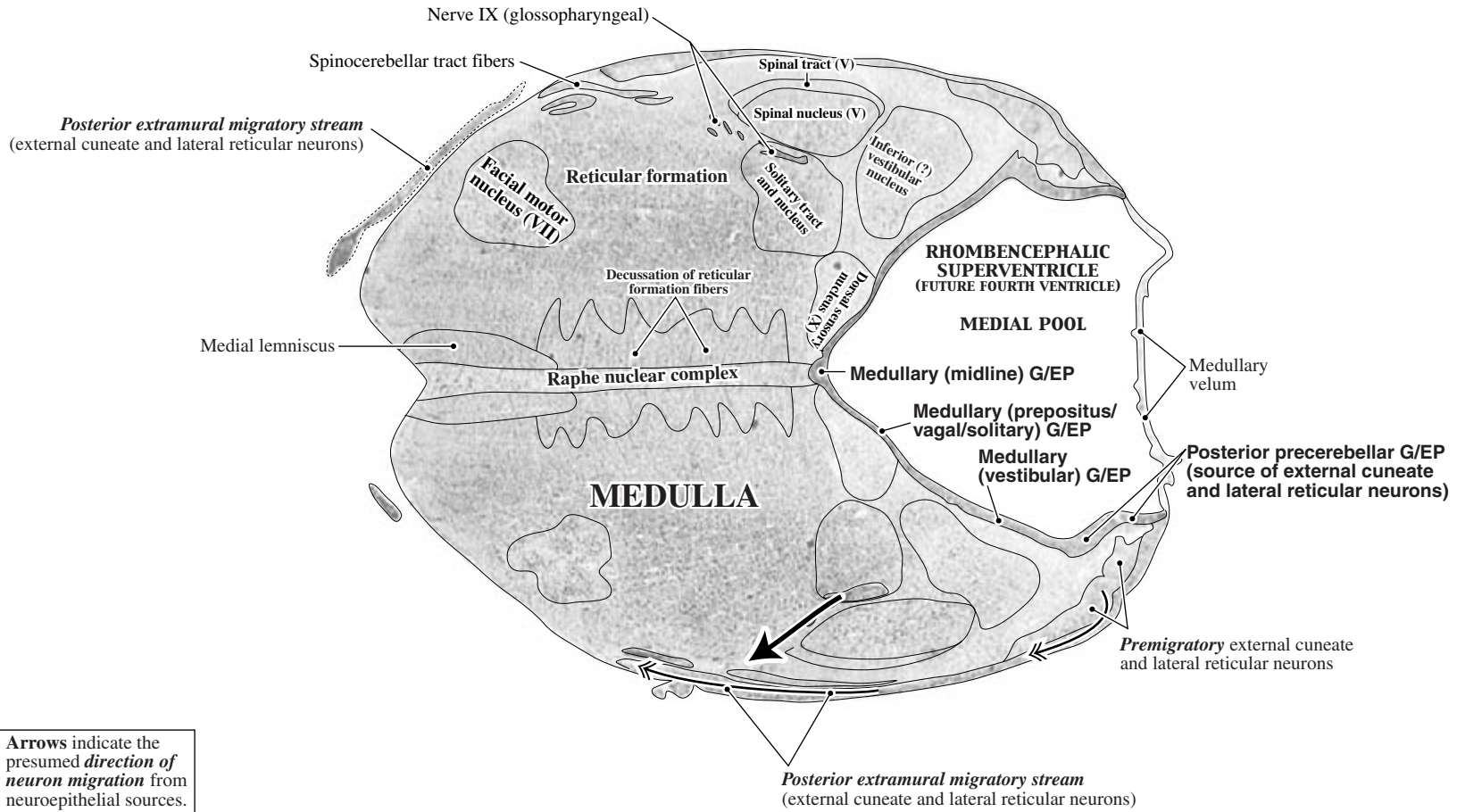


Levels 14 to 21 are only shown at high magnification.

PLATE 53B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioeptihelium/ependyma



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in nerves and fiber tracts.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 54A

**GW11 Horizontal
CR 57 mm, C1500
Level 19: Section 2202**

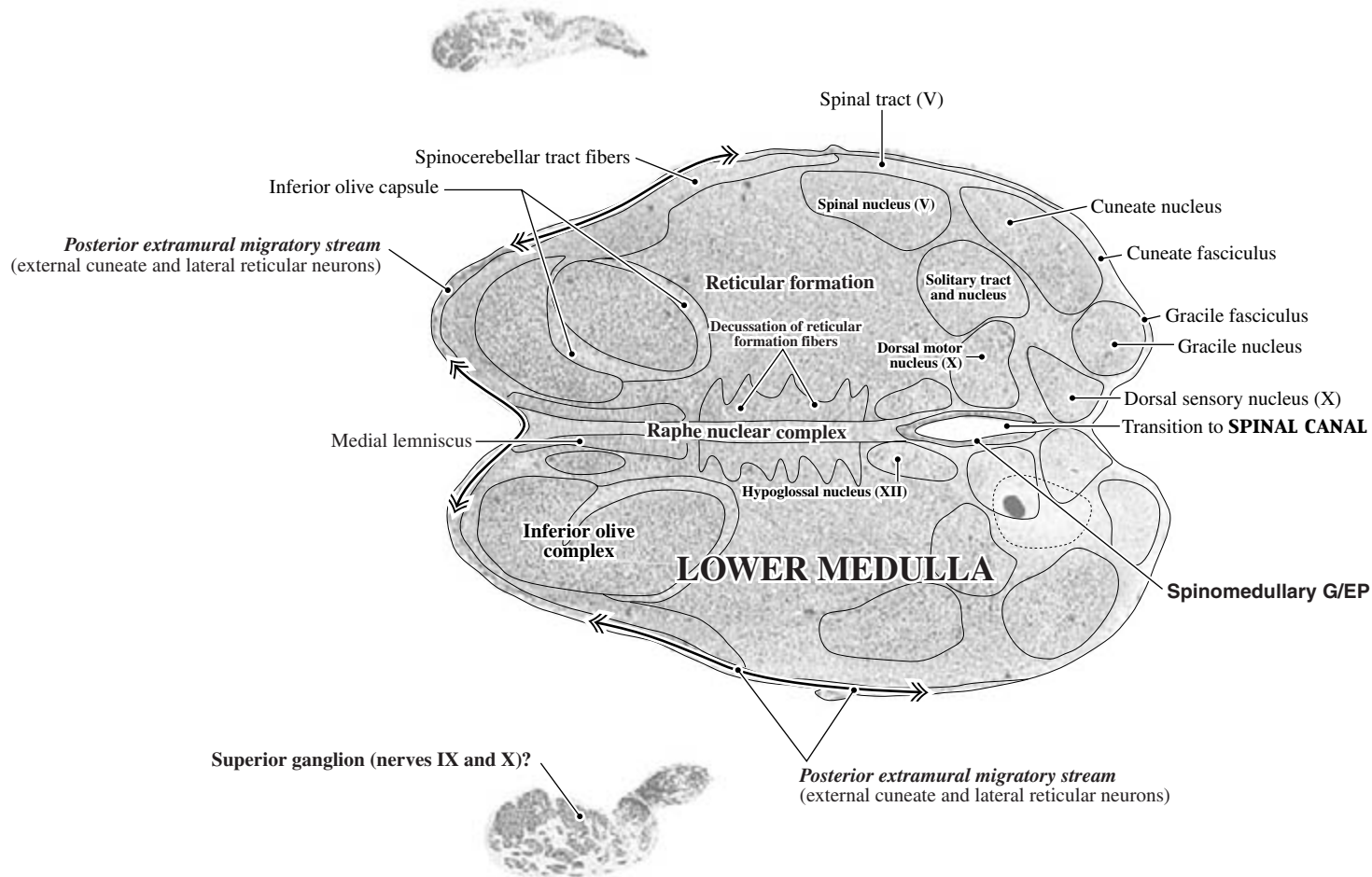


Levels 14 to 21 are only shown at high magnification.

PLATE 54B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioepithelium/ependyma



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Dashed lines indicate staining and/or sectioning artifacts.

PLATE 55A

**GW11 Horizontal
CR 57 mm, C1500
Level 20: Section 2322**

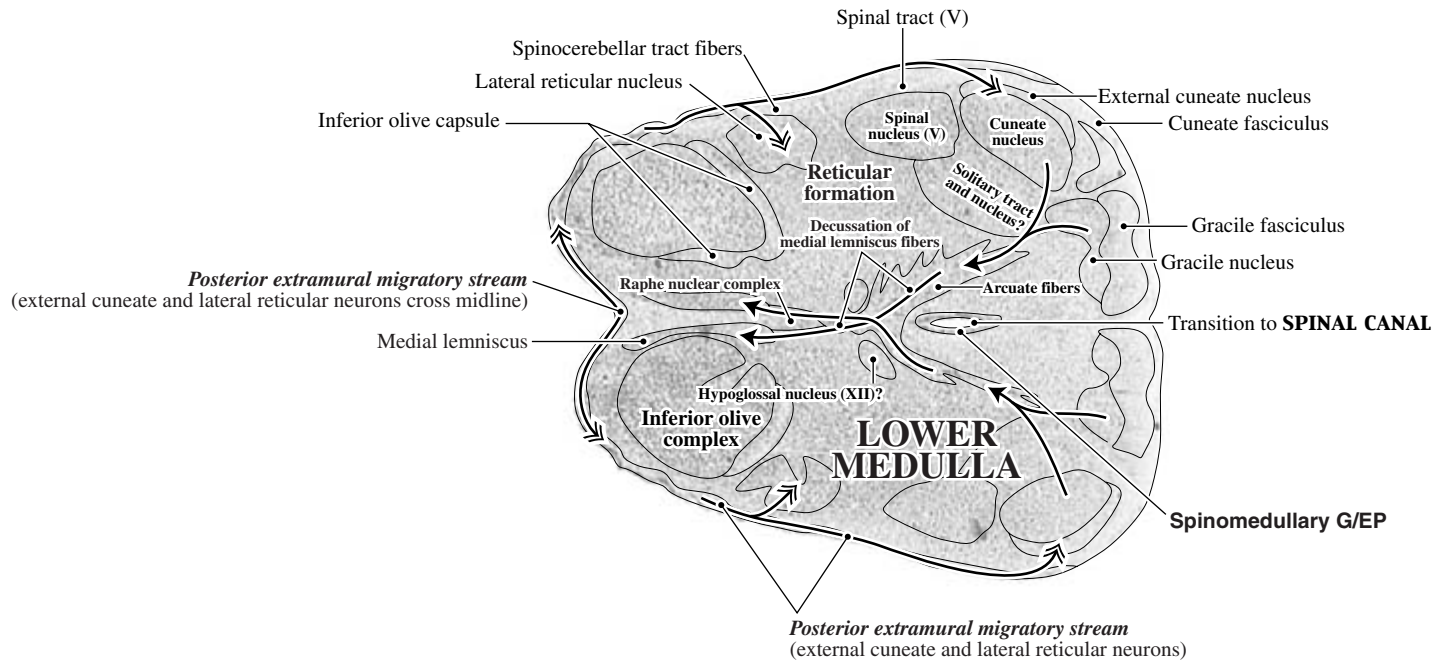


Levels 14 to 21 are only shown at high magnification.

PLATE 55B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioepithelium/ependyma



Superior ganglion (nerves IX and X)?

Inferior ganglion (nerves IX and X)?

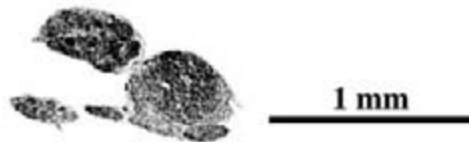
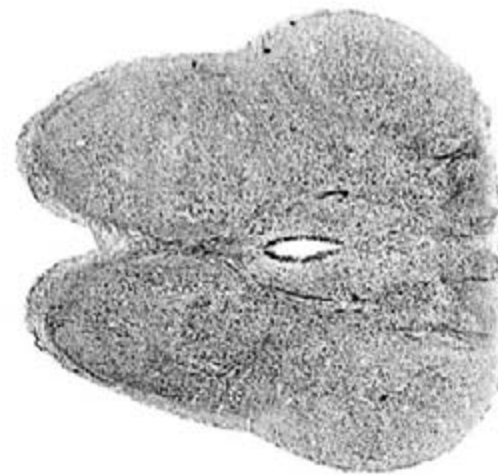


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 56A

**GW11 Horizontal
CR 57 mm, C1500
Level 21: Section 2402**

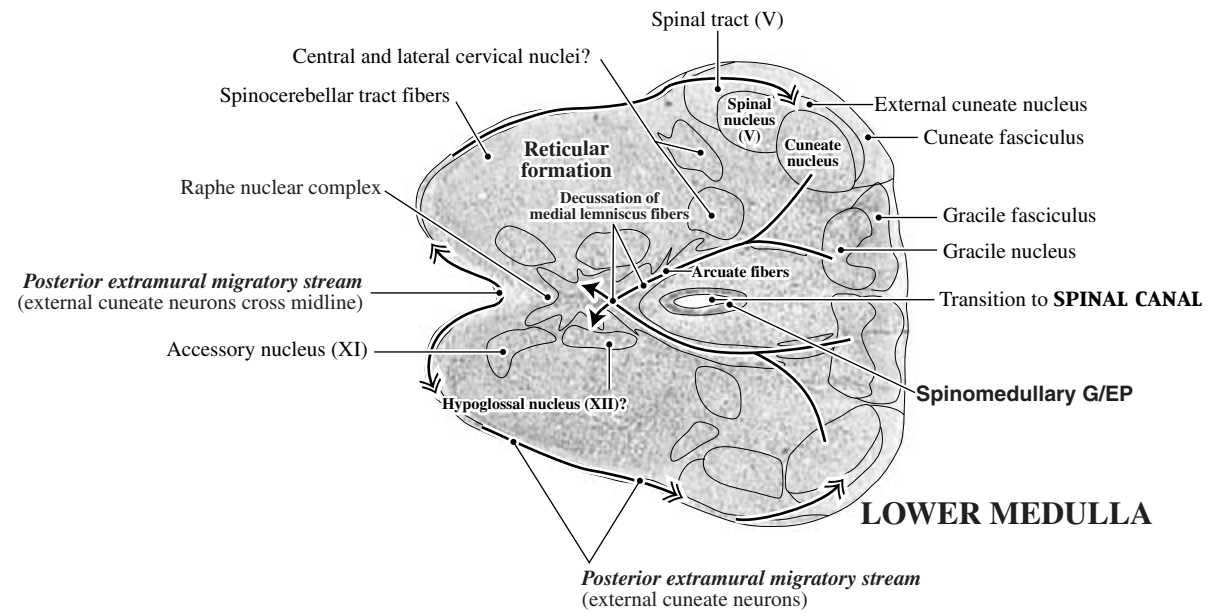


Levels 14 to 21 are only shown at high magnification.

PLATE 56B

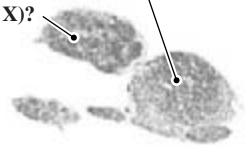
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

G/EP - Glioepithelium/ependyma



Superior ganglion (nerves IX and X)?

Inferior ganglion (nerves IX and X)?



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PART IV: GW9 SAGITTAL

This is specimen number 6658 in the Carnegie Collection, designated here as C6658, a female with a crown-rump length (CR) of 40 mm estimated to be at gestational week (GW) 9. The entire fetus was cut in the sagittal plane in 40 μ m sections and stained with hematoxylin and eosin. Information on the date of specimen collection, fixative, and embedding medium (appears to be celloidin) was not available to us. The histology is excellent, and this is one of the best preserved specimens in any of the Collections at the National Museum of Health and Medicine. Since there is no photograph of this specimen's brain before histological processing, a specimen from Hochstetter (1919) that is comparable in age to C6658 is used to show external brain features at GW9 (**A, Figure 3**). C6658's brain structures are more difficult to understand because the sections are not cut parallel to the midline; **Figure 3** shows the approximate rotations in horizontal (**B**) and vertical (**C**) dimensions. Photographs of 11 sections (**Levels 1-11**) are illustrated at low magnification in four-parts (**Plates 57A-D** through **67A-D**). The **A/B** parts show the brain in place in the skull; the **C/D** parts show only the brain (and some peripheral ganglia) at slightly higher magnification. **Plates 68-83** show high-magnification views of various parts of the brain at different levels from the cerebral cortex (**Plate 68**) to the pons and sensory ganglia (**Plates 82-83**). All of the high-magnification plates are rotated 90° (landscape orientation) to more efficiently use page space.

C6658 is considerably less mature than the GW11 specimens. Throughout the cerebral cortex, the *neuroepithelium* is prominent and appears to be without a subventricular zone. The *stratified transitional field (STF)* contains *STF1* and *STF5* throughout; with *STF4* only in lateral areas. The most prominent developmental feature of the cerebral cortex is that both the *STF* layers and the cortical plate have a pronounced anterolateral (thicker) to dorso-medial (thinner) maturation gradient. The olfactory bulb is just beginning to evaginate in front of the basal telencephalic neuroepithelium. In anterolateral parts of the cerebral cortex, streams of neurons and glia appear to leave *STF4* and enter the *lateral migratory stream*. The hip-

pocampus contains *ammonic and dentate migrations*, but there is no evidence of a pyramidal in Ammon's horn or a dentate gyrus. A massive *neuroepithelium/subventricular zone* overlies the amygdala, nucleus accumbens, and striatum (caudate and putamen) where neurons (and glia) are being generated.

The cerebellum is a thick, smooth plate overlying the posterior pons and medulla, and a definite *neuroepithelium* at the ventricular surface, indicating some Purkinje cells are still being generated. Many Purkinje cells are sojourning in a dense layer outside the neuroepithelium, and others are migrating upward. Many of the deep neurons are superficial in the cerebellum, but some are migrating downward to intermingle with upwardly migrating Purkinje cells. The cortical surface is partially covered by an *external germinal layer (egl)* that is actively producing neuronal stem cells, as it grows over the surface of the cerebellar cortex.

The third ventricle, aqueduct, and fourth ventricle are lined by thin *neuroepithelia*. The midbrain tegmentum, pons, and medulla have the thinnest neuroepithelia indicating that only the latest generated neurons are being produced at this time. The thick precerebellar neuroepithelium is an exception in the medulla. Thicker neuroepithelia are in the cerebellum (see above) and midbrain tectum, indicating many neurons are still being generated, although the majority of the neurons in these sites are already postmitotic. The neuroepithelium is still thicker in the hypothalamus and thalamus, in accordance with the later maturation of the diencephalon compared to the rest of the brainstem.

Neurons throughout the diencephalon, midbrain tegmentum, pons, and medulla are migrating and settling. Nuclear divisions are very indistinct throughout the diencephalon. More definition is seen in the midbrain tegmentum, pons, and medulla. The large *anterior extramural, posterior extramural, and intramural migratory streams* are prominent in the medulla and pons.

GW9 SAGITTAL

A perfect sagittal cut through the brain bisects the cerebral cortex into two separate hemispheres by passing through the interhemispheric fissure, and does the same in the brainstem by passing through the midline of the ventricles.

Sections of C6658's brain are not parallel to the midline either horizontally (-11.71° , top view) or vertically (-6.64° , back view). In each of the illustrated sections on the following pages, the anterior edge of the cortex (top right) is tilted away from the observer, while the medulla and upper spinal cord (bottom) are tilted toward the observer.

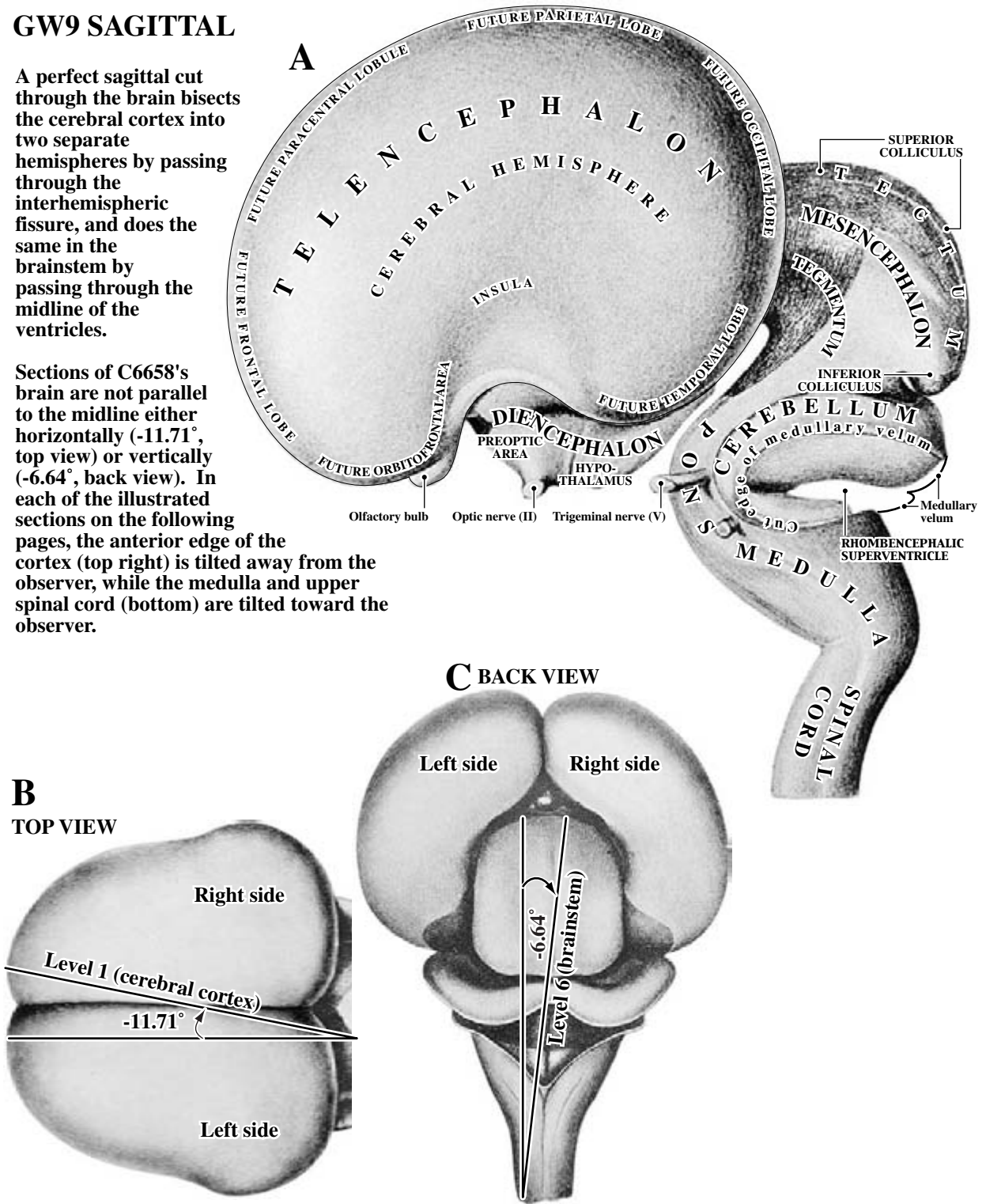
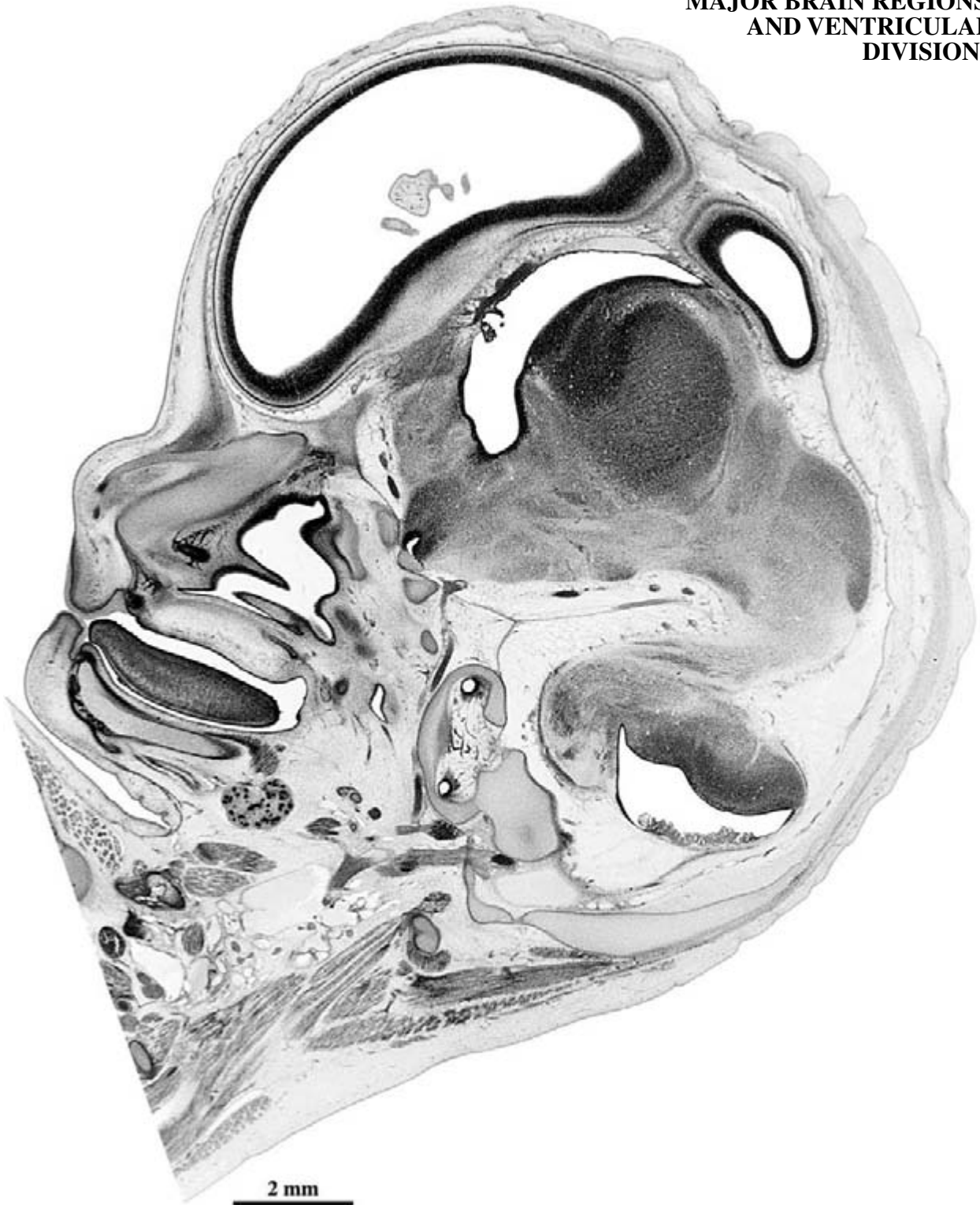
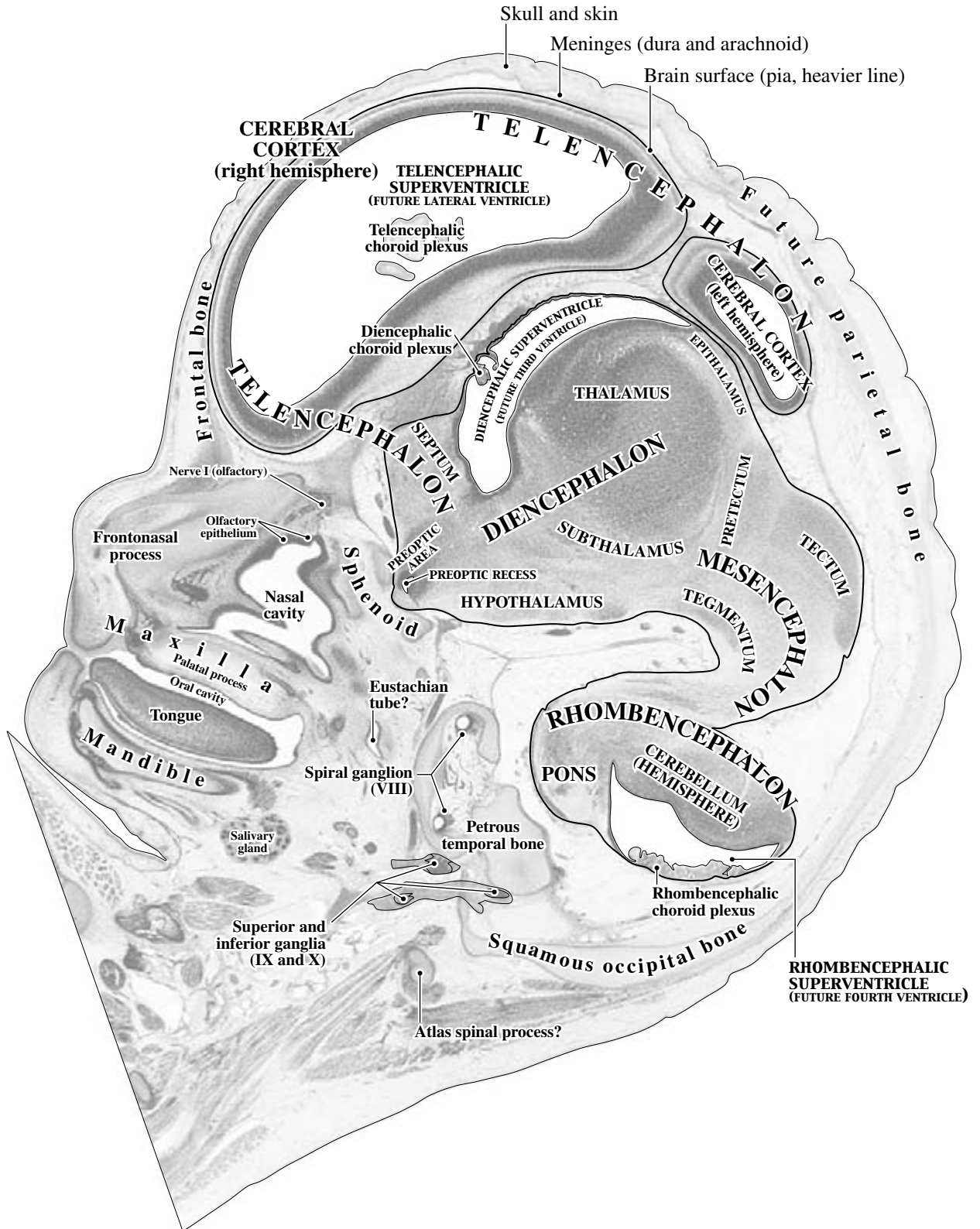


Figure 3. A, Lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 38 mm (modified from Figure 43, Table VII, Hochstetter, 1919) identifies external features of a brain similar to C6658 (CR 40 mm). B, Top view of the brain in A (modified from Figure 45, Table VIII, Hochstetter, 1919) shows how C6658's sections rotate from a line parallel to the horizontal midline in the interhemispheric fissure. C, Back view of the brain in A (modified from Figure 44, Table VIII, Hochstetter, 1919) shows how C6658's sections rotate from a line parallel to the vertical midline in the brainstem and upper cervical spinal cord.

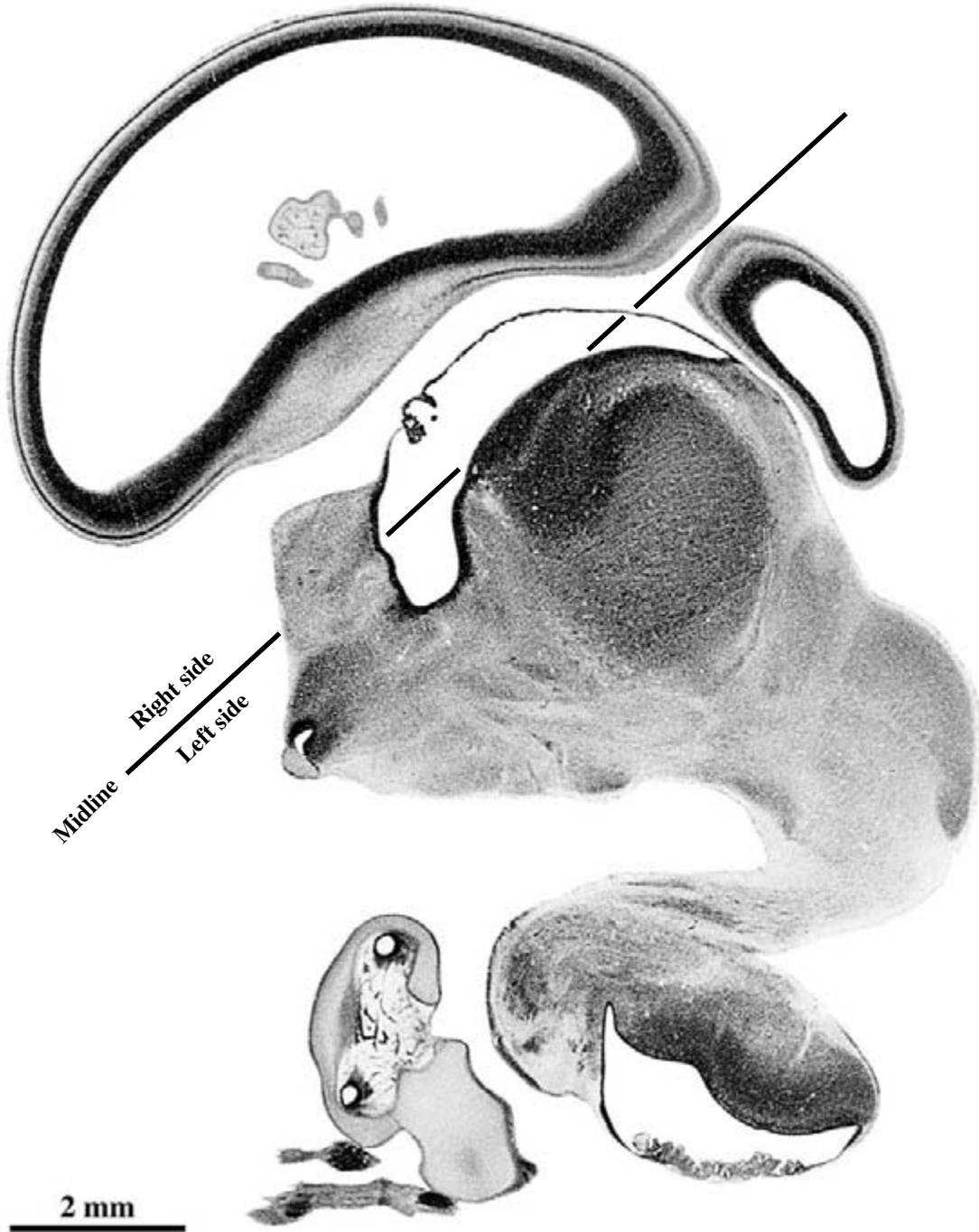


Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

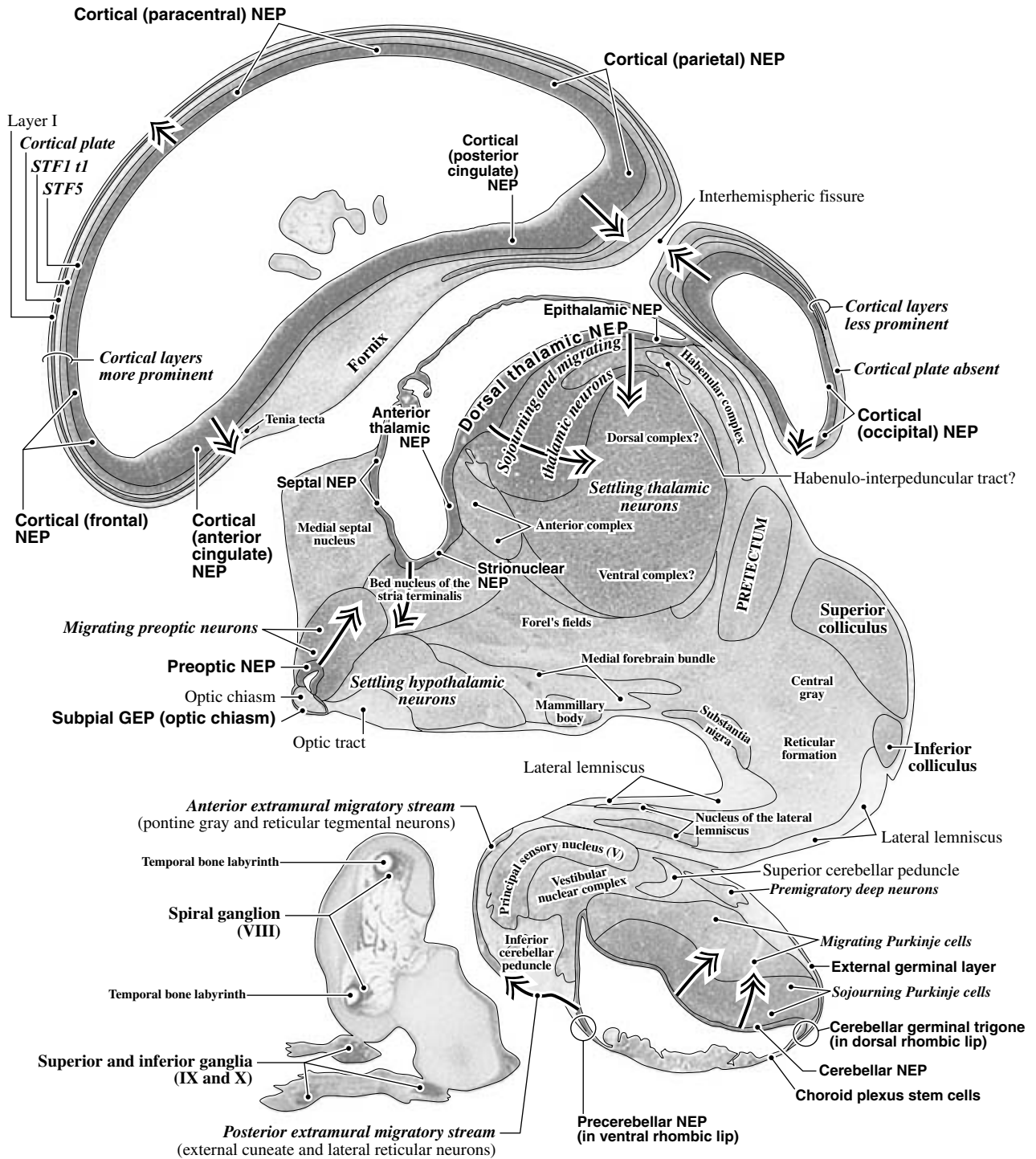
PLATE 57B



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

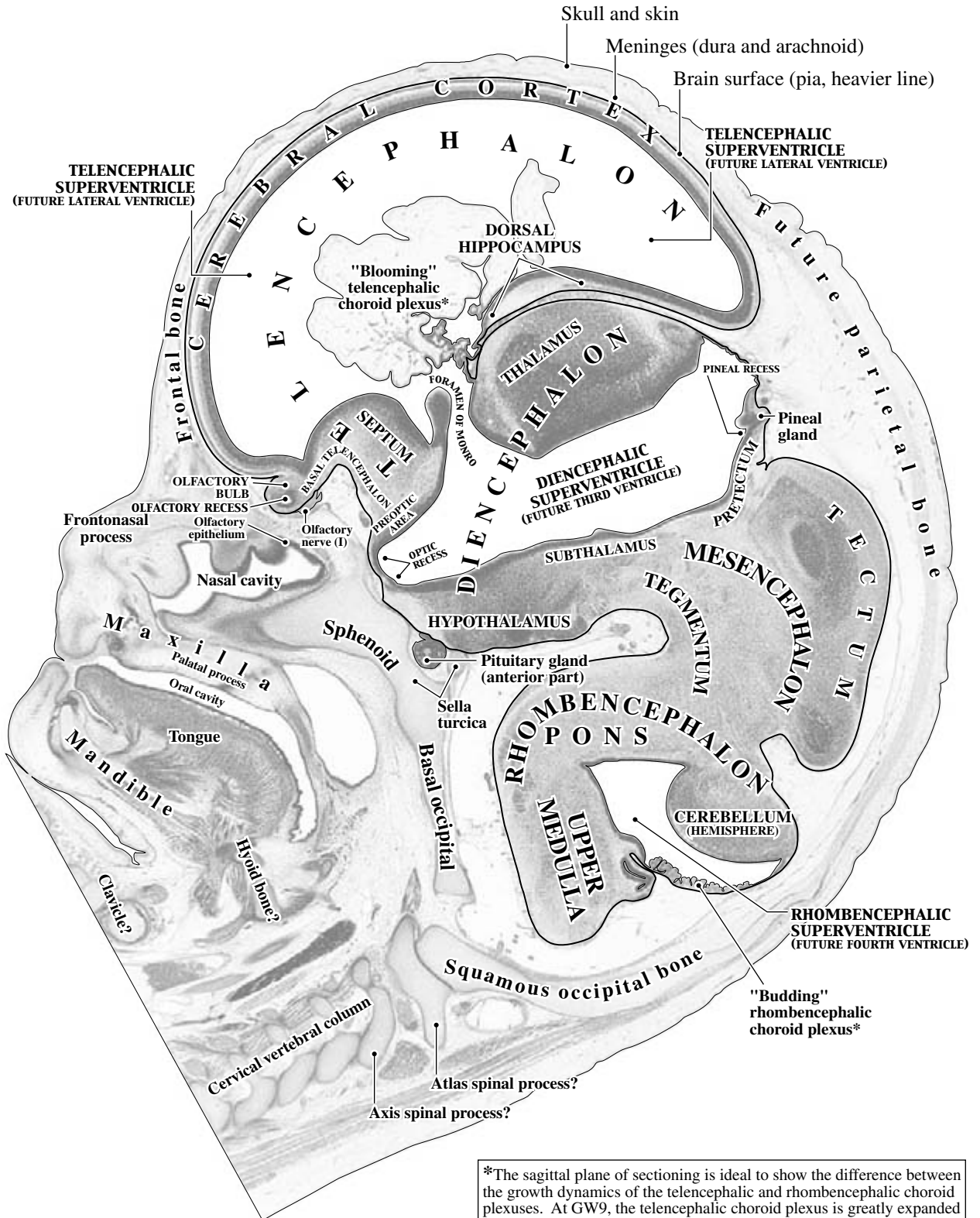
ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.



Neuroepithelial divisions, glioeptithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 58B

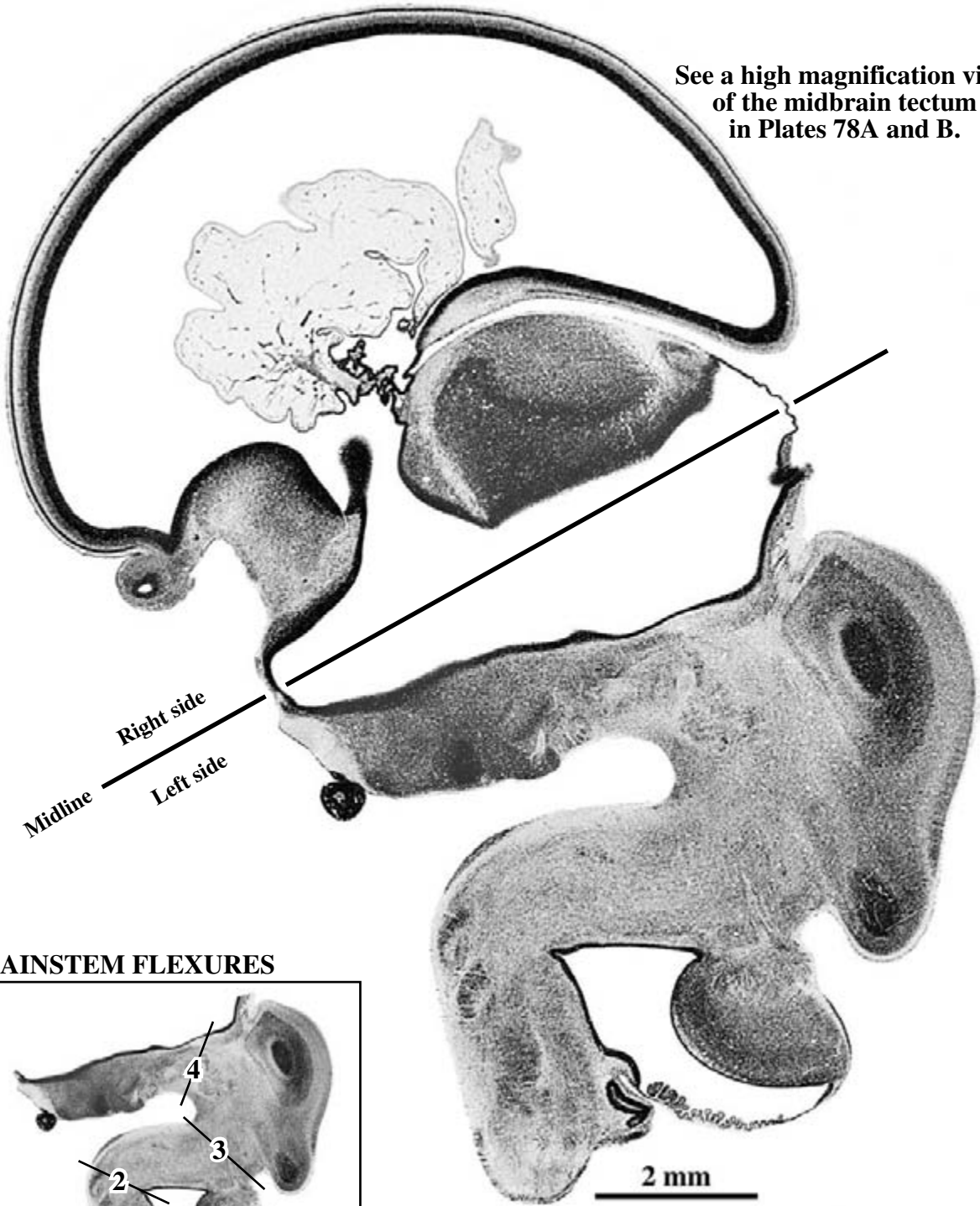


*The sagittal plane of sectioning is ideal to show the difference between the growth dynamics of the telencephalic and rhombencephalic choroid plexuses. At GW9, the telencephalic choroid plexus is greatly expanded ("blooming") but the rhombencephalic choroid plexus is still small ("budding"). The rhombencephalic choroid plexus "blooms" during the second and third trimesters (see Volumes 3 and 2).

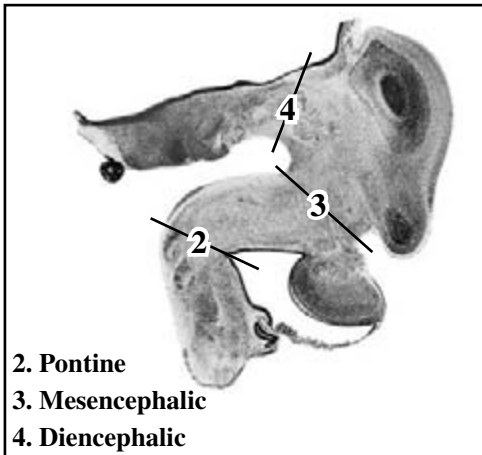
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

GW9 Sagittal
CR 40 mm, C6658
Level 2: Slide 63, Section 1
NEUROEPITHELIAL/GLIOEPITHELIAL DIVISIONS AND
DIFFERENTIATING BRAIN STRUCTURES

See a high magnification view
of the midbrain tectum
in Plates 78A and B.

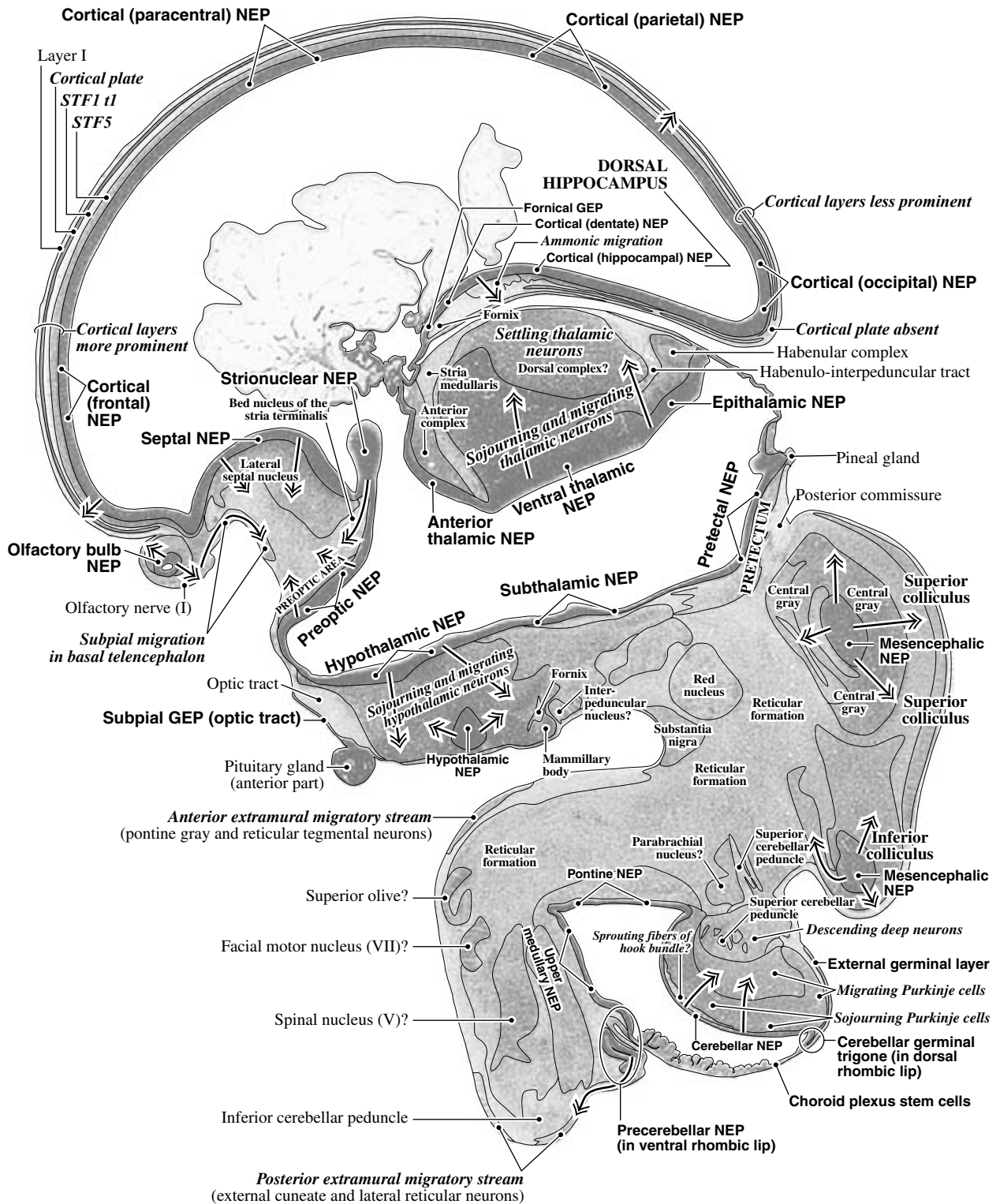


BRAINSTEM FLEXURES



- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

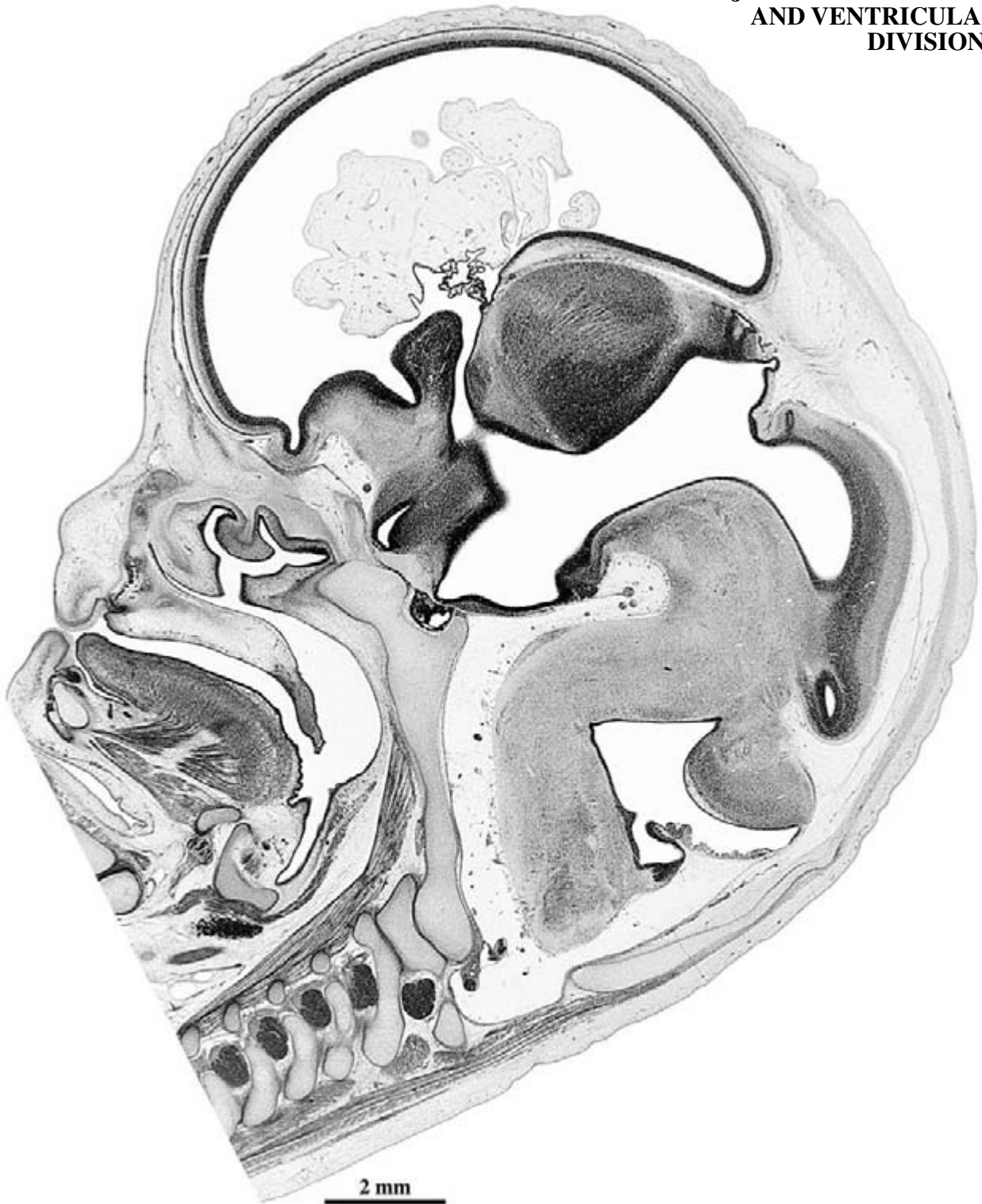
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

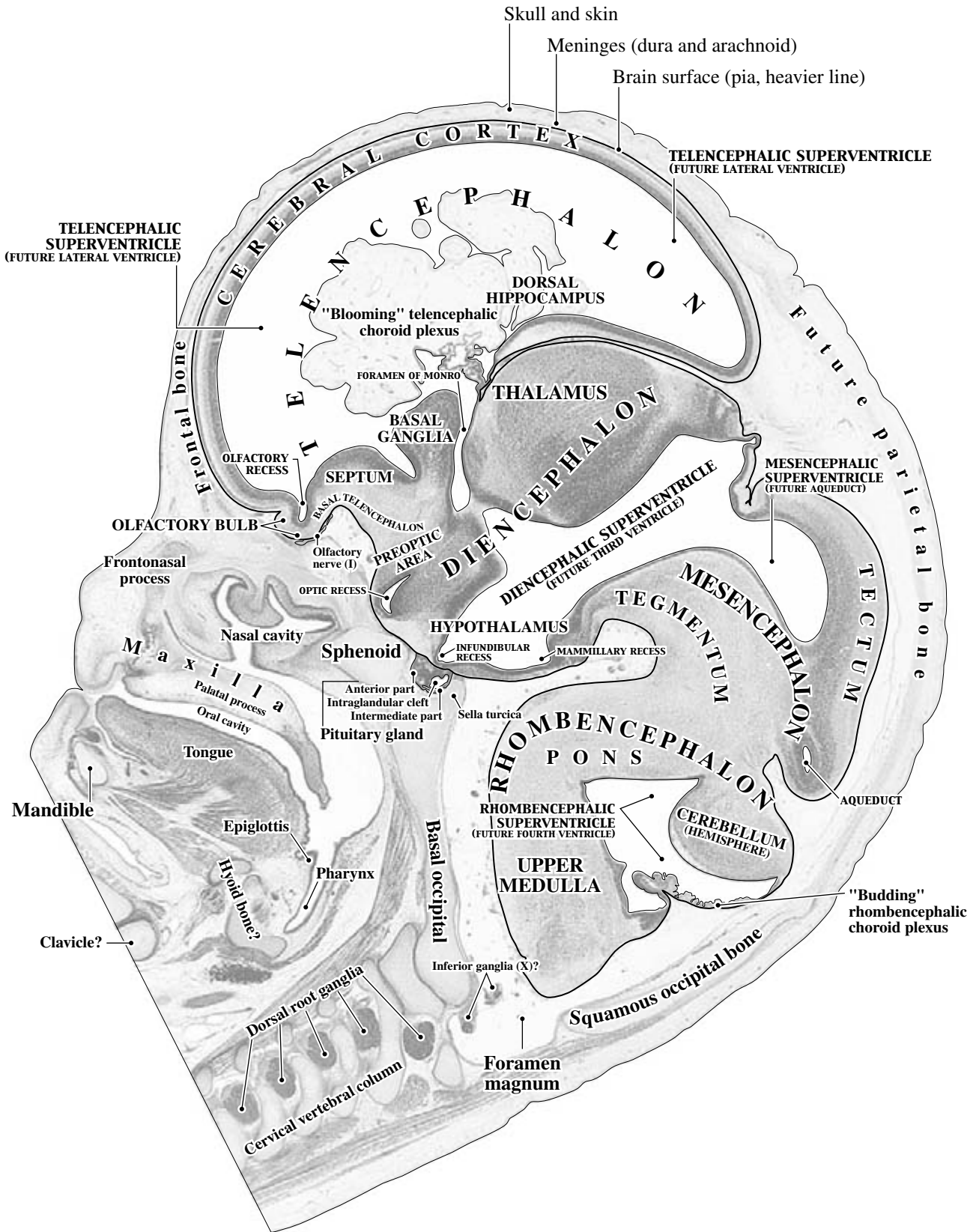
ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

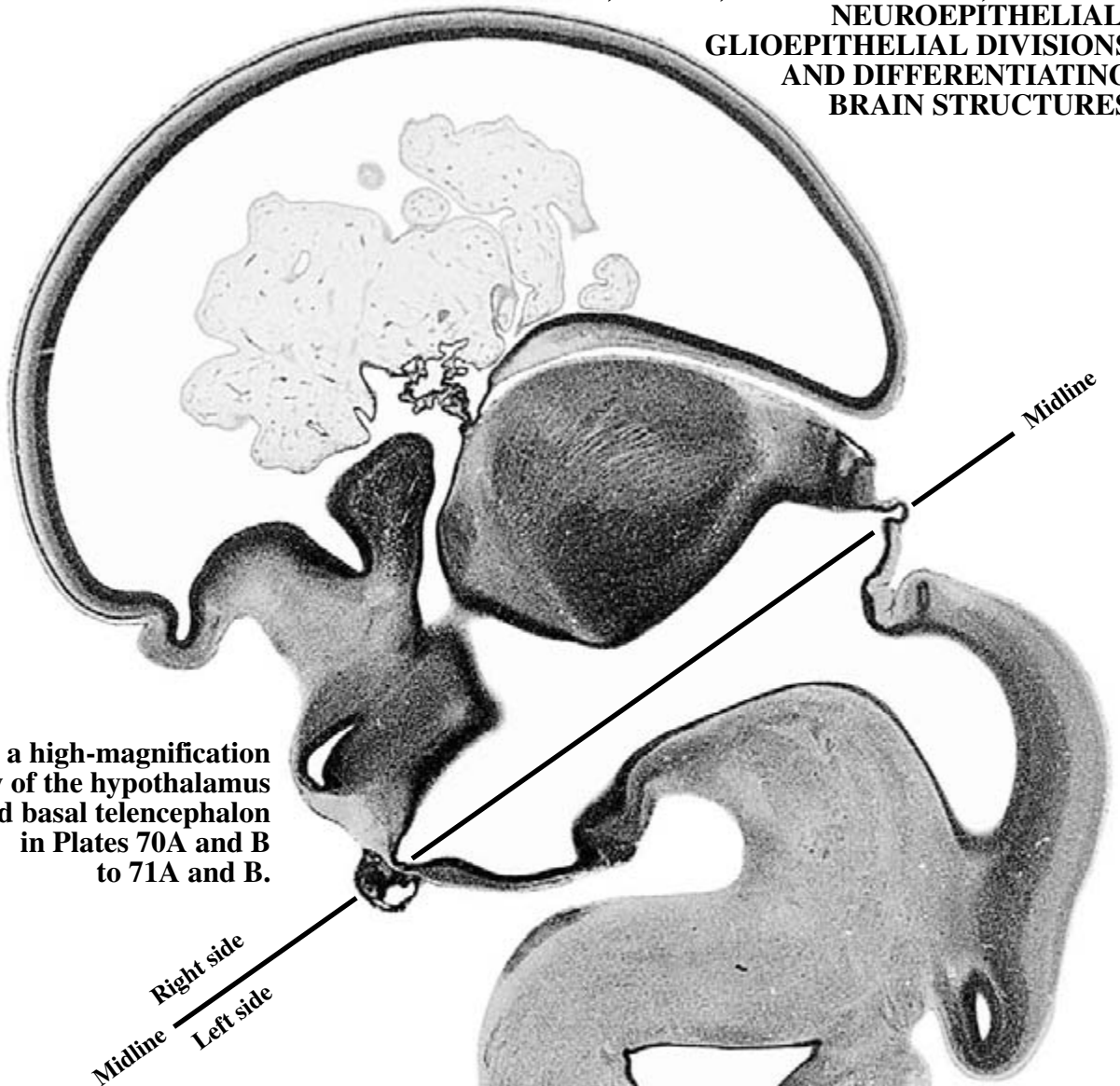


Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 59B



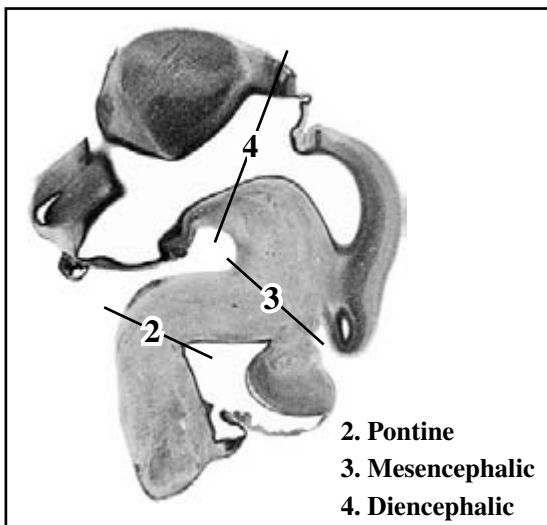
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See a high-magnification view of the hypothalamus and basal telencephalon in Plates 70A and B to 71A and B.

Right side
Left side
Midline

BRAINSTEM FLEXURES

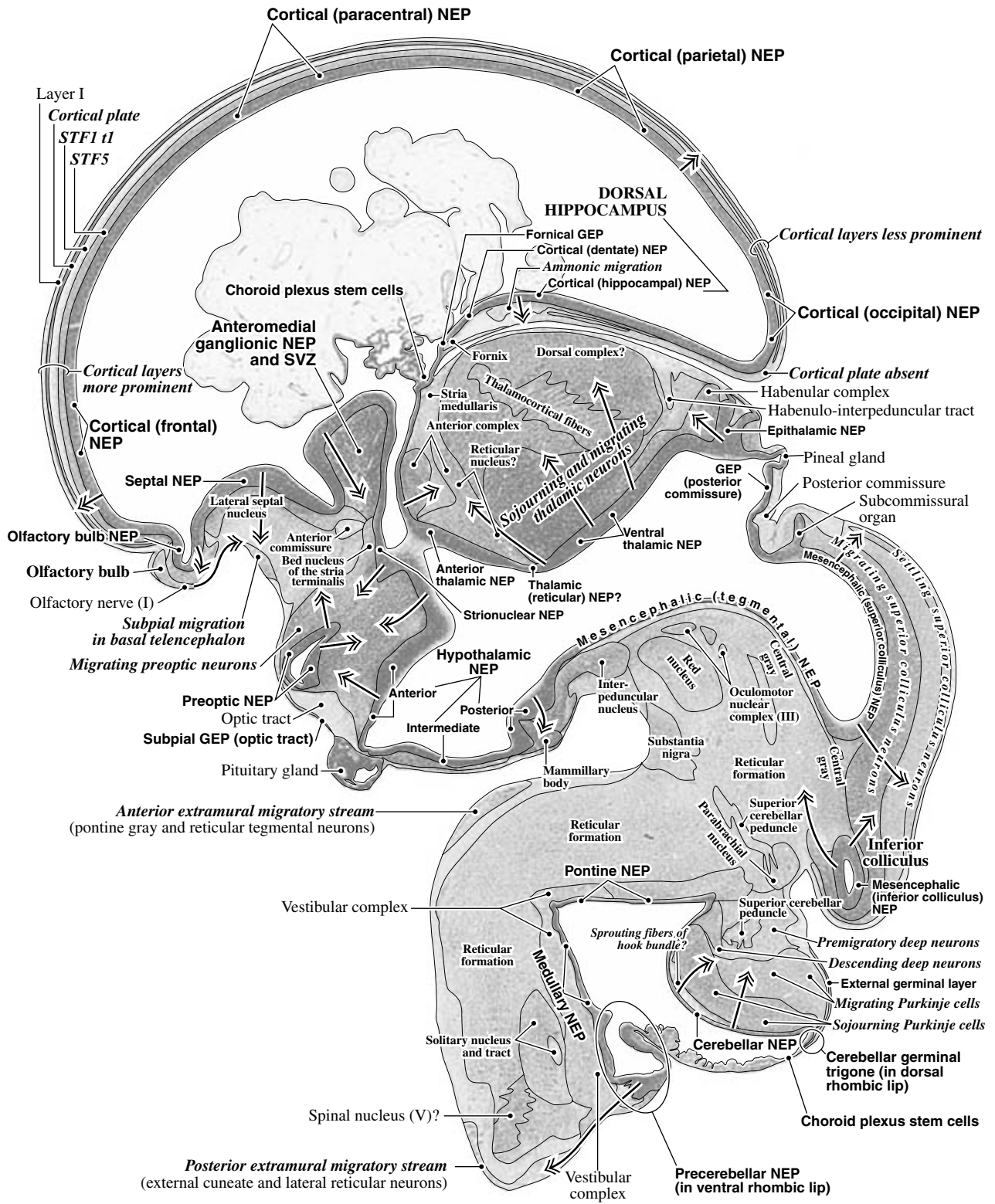


- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

See a high-magnification view of the midbrain in Plates 77A and B.

2 mm

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

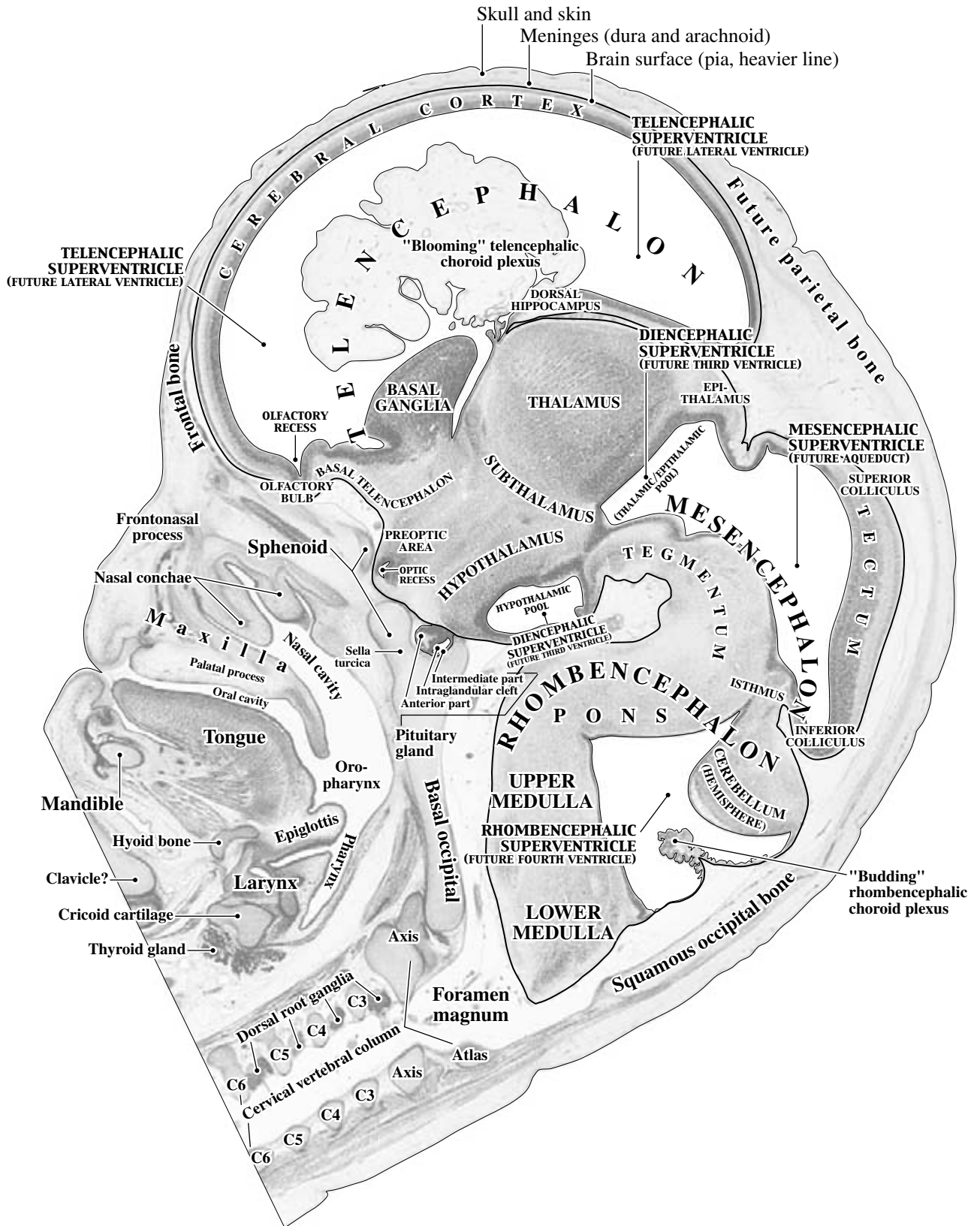
ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field
 SVZ - Subventricular zone

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.



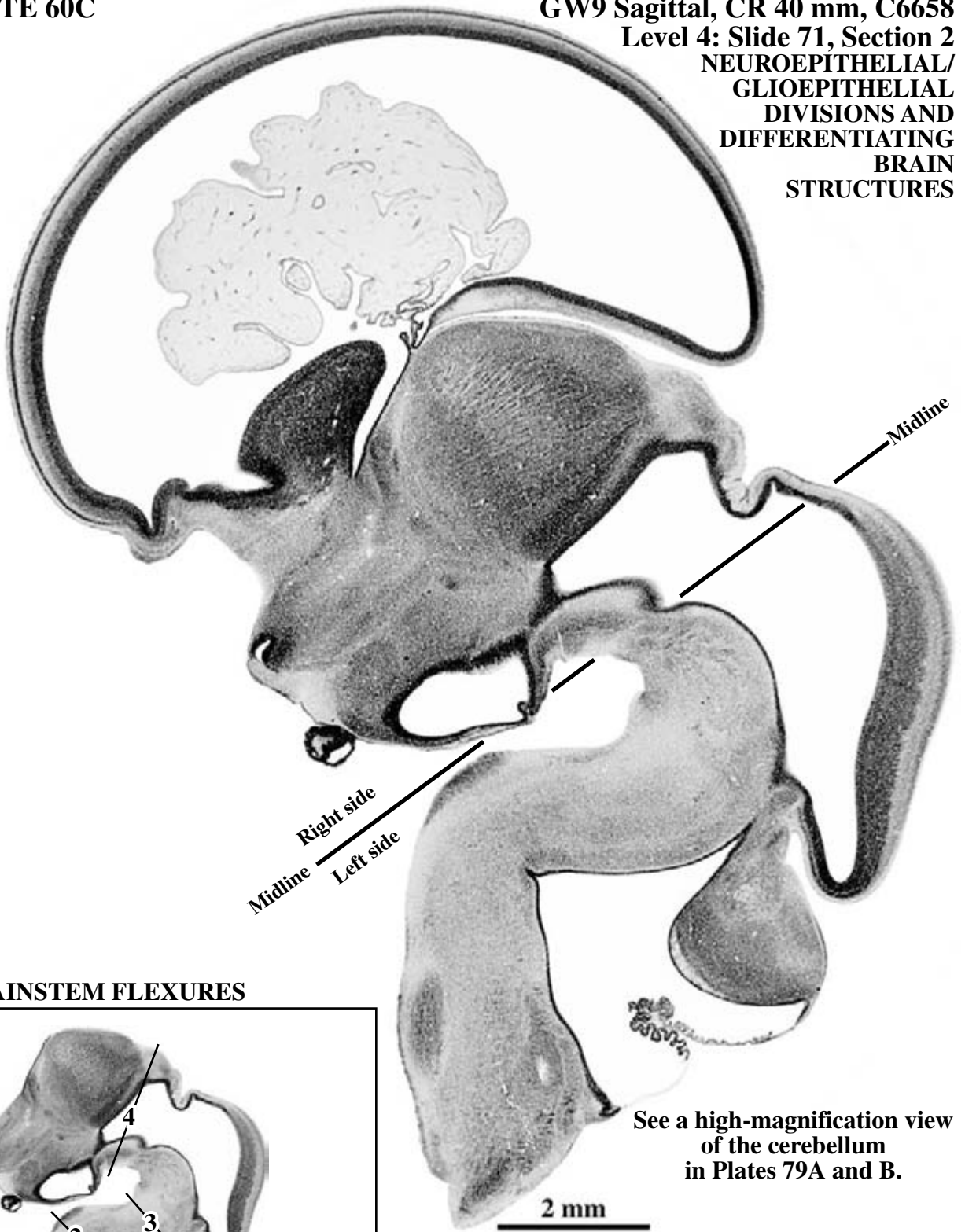
Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 60B

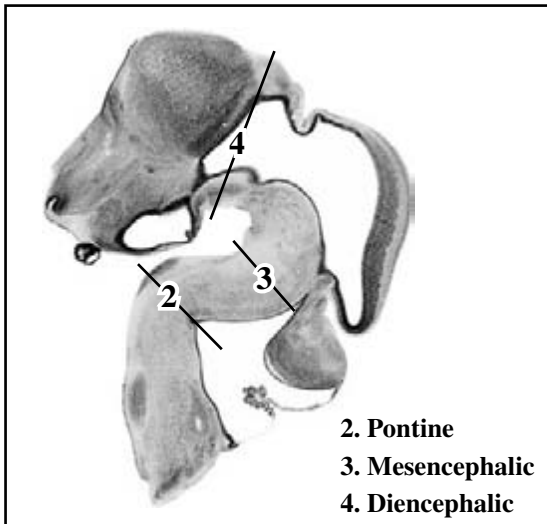


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

GW9 Sagittal, CR 40 mm, C6658
Level 4: Slide 71, Section 2
NEUROEPITHELIAL/
GLIOEPITHELIAL
DIVISIONS AND
DIFFERENTIATING
BRAIN
STRUCTURES

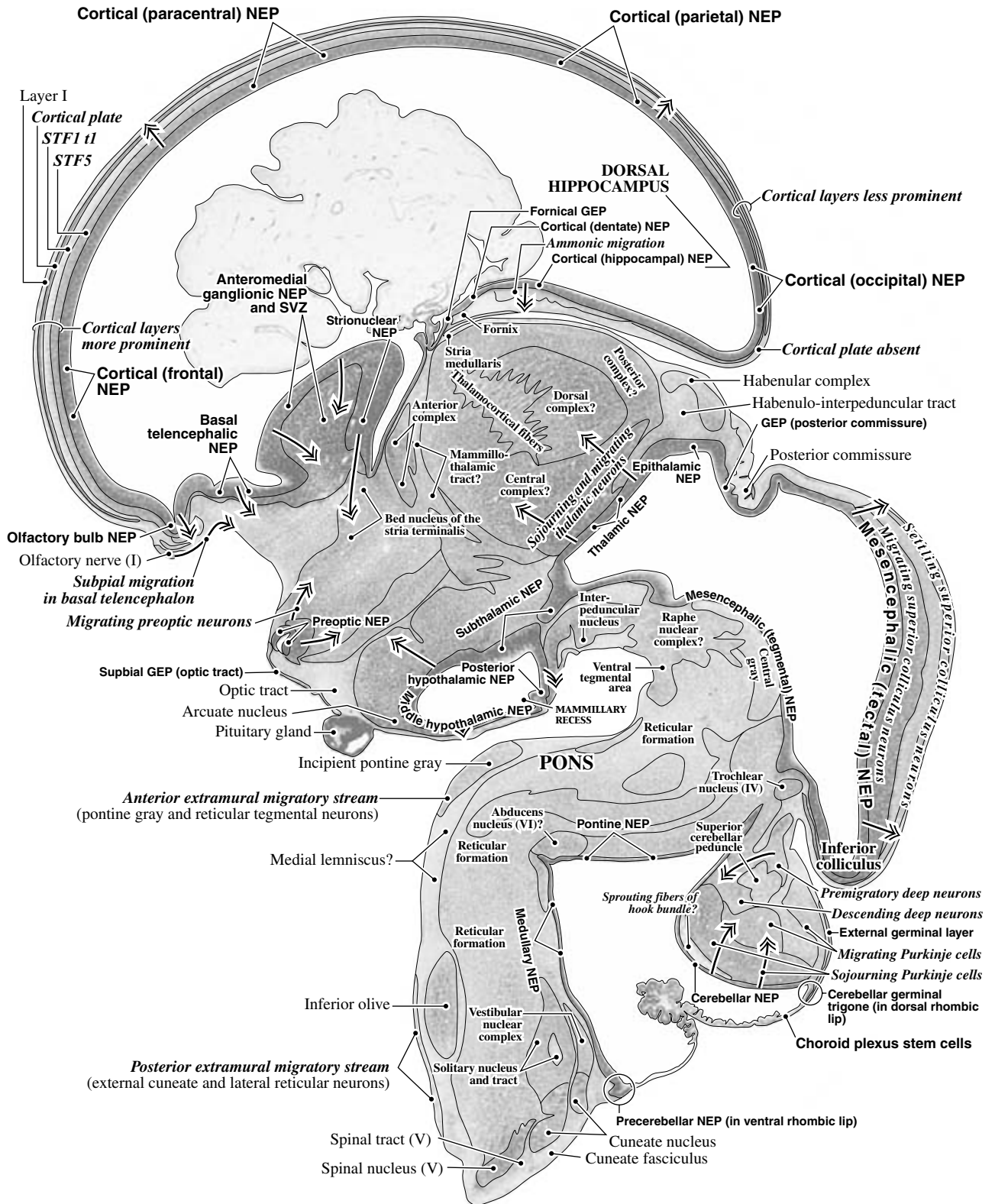


BRAINSTEM FLEXURES



See a high-magnification view of the cerebellum in Plates 79A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

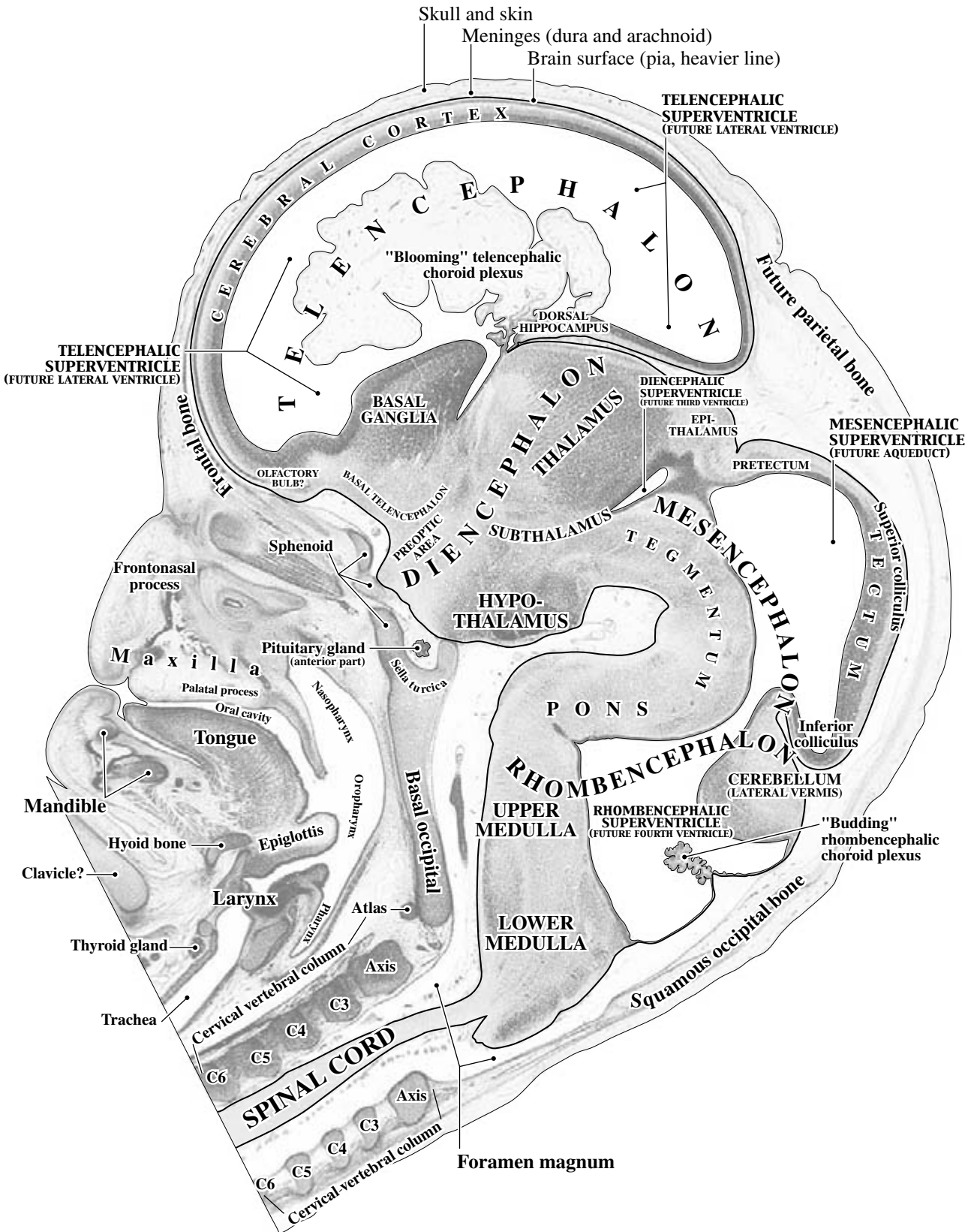
ABBREVIATIONS:
 GEP - Gliopithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field
 SVZ - Subventricular zone

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.



Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

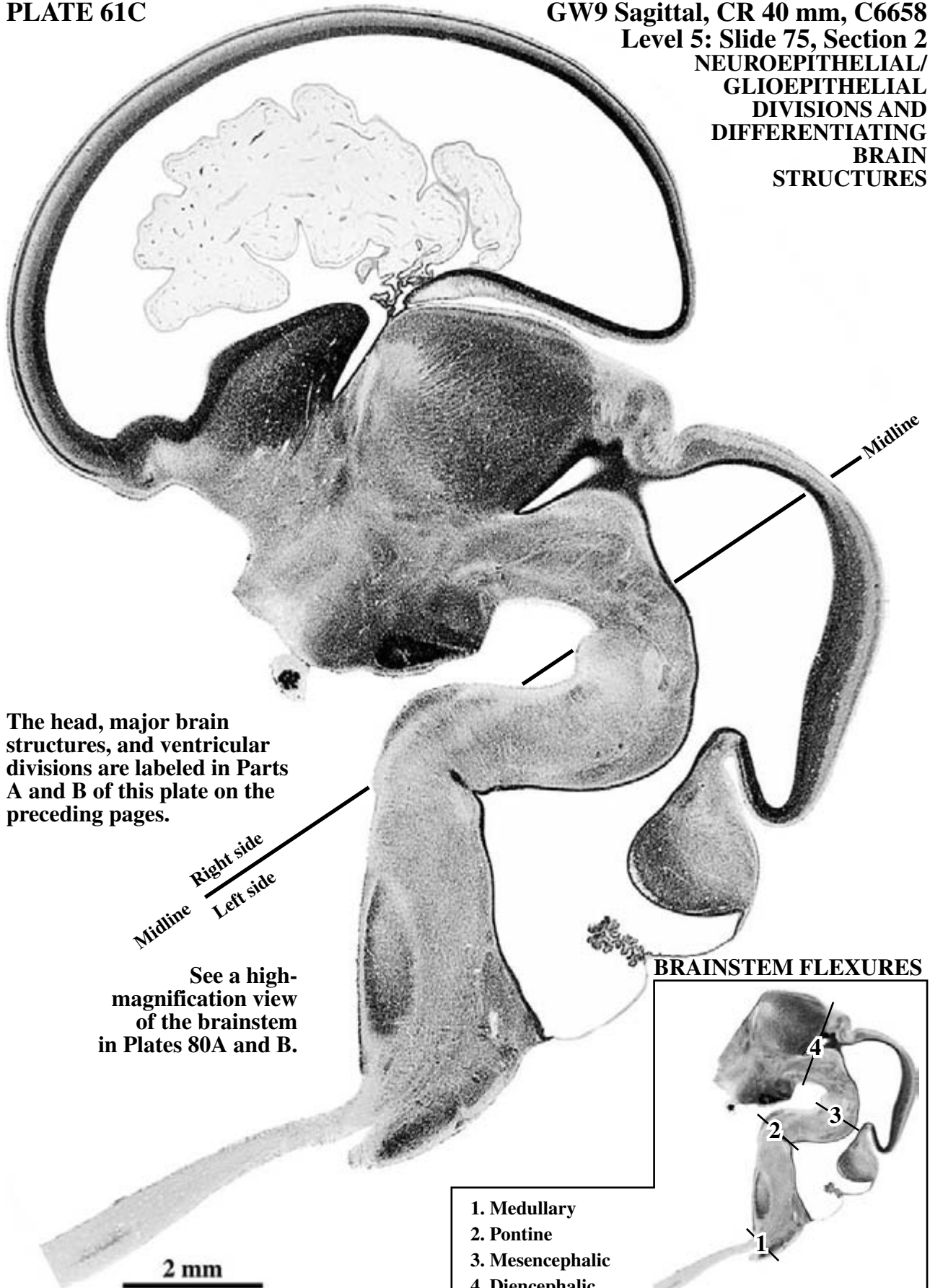
PLATE 61B



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

PLATE 61C

GW9 Sagittal, CR 40 mm, C6658
Level 5: Slide 75, Section 2
NEUROEPITHELIAL/
GLIOEPITHELIAL
DIVISIONS AND
DIFFERENTIATING
BRAIN
STRUCTURES



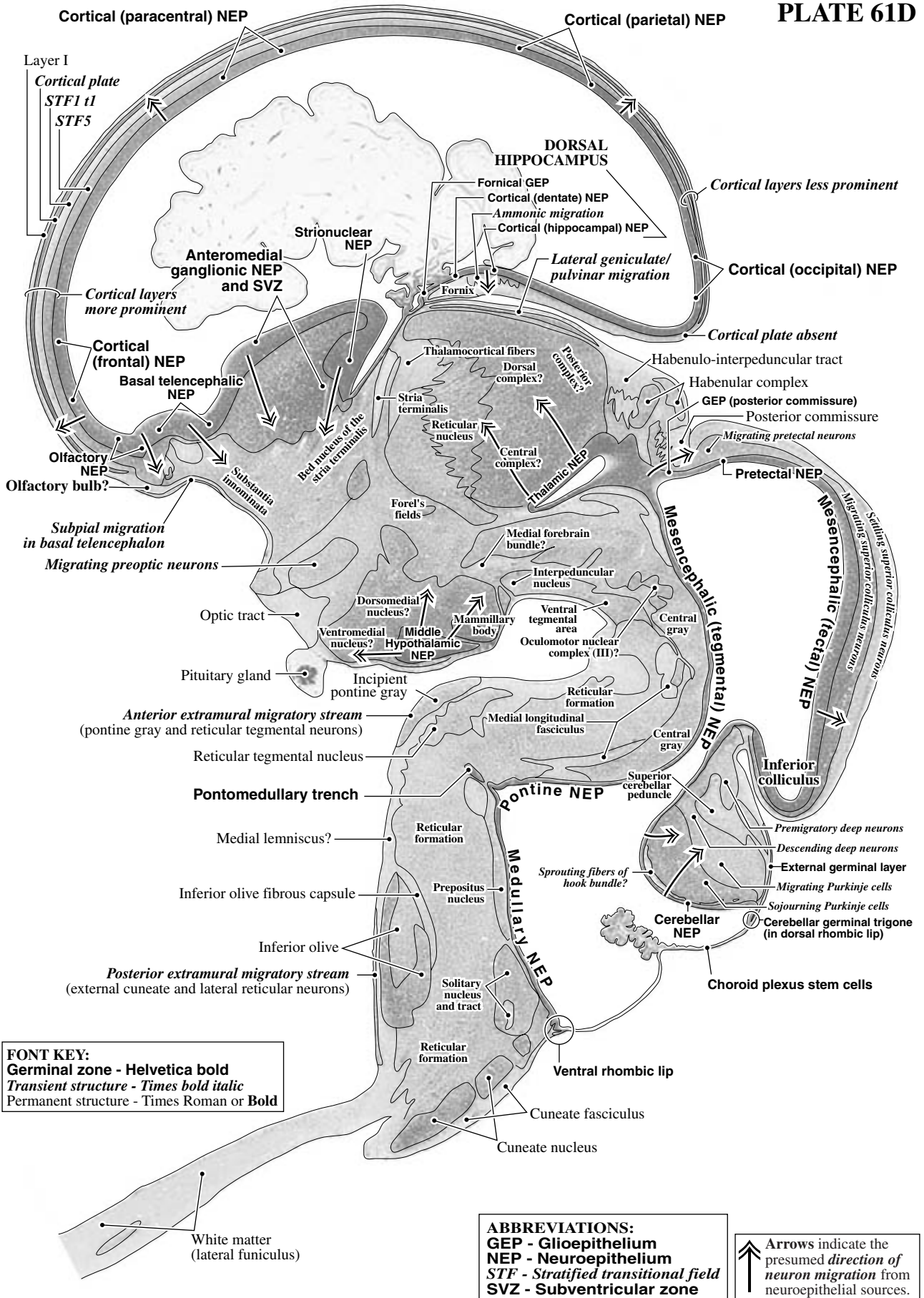
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

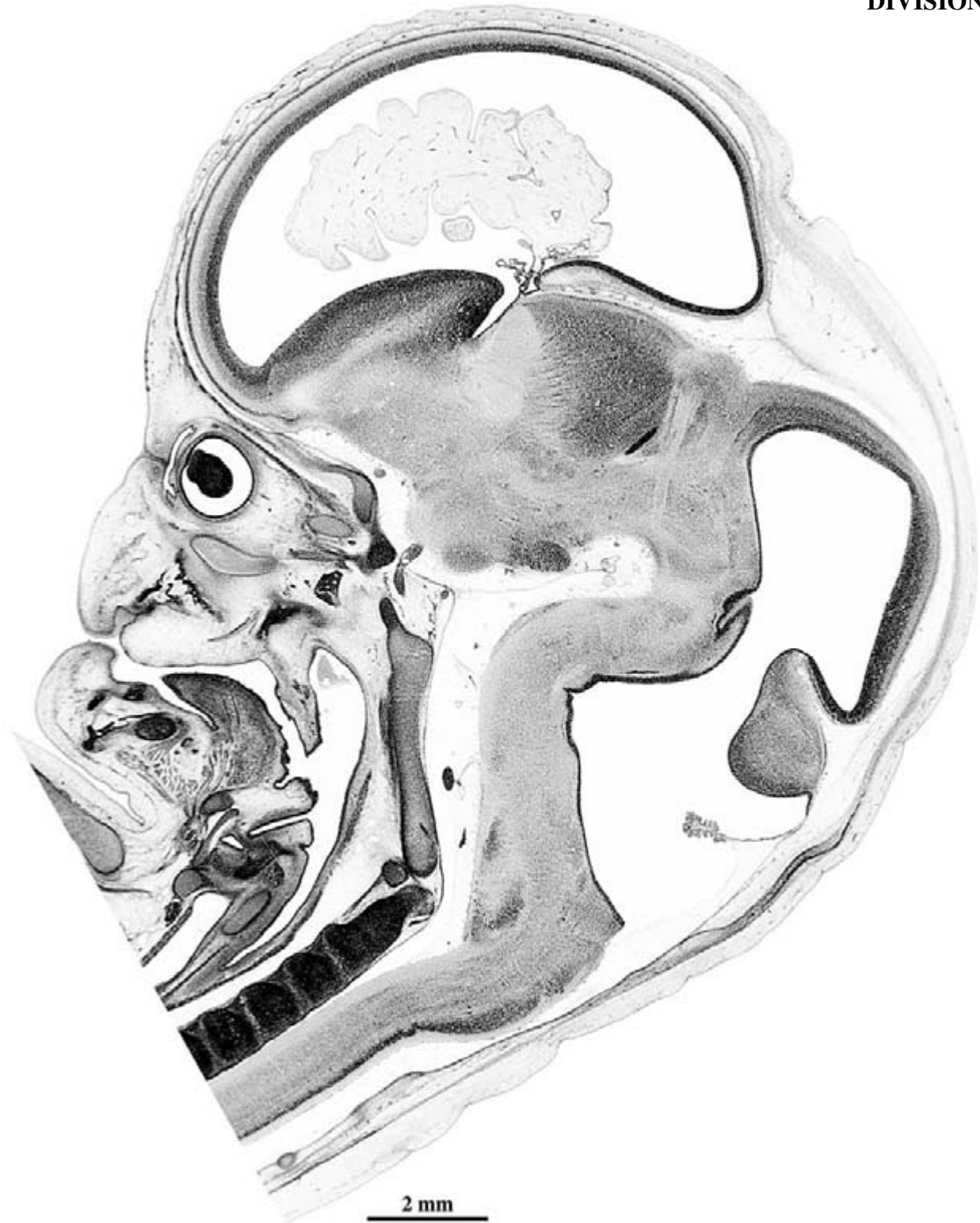
Midline
Right side
Left side

See a high-magnification view of the brainstem in Plates 80A and B.

BRAINSTEM FLEXURES

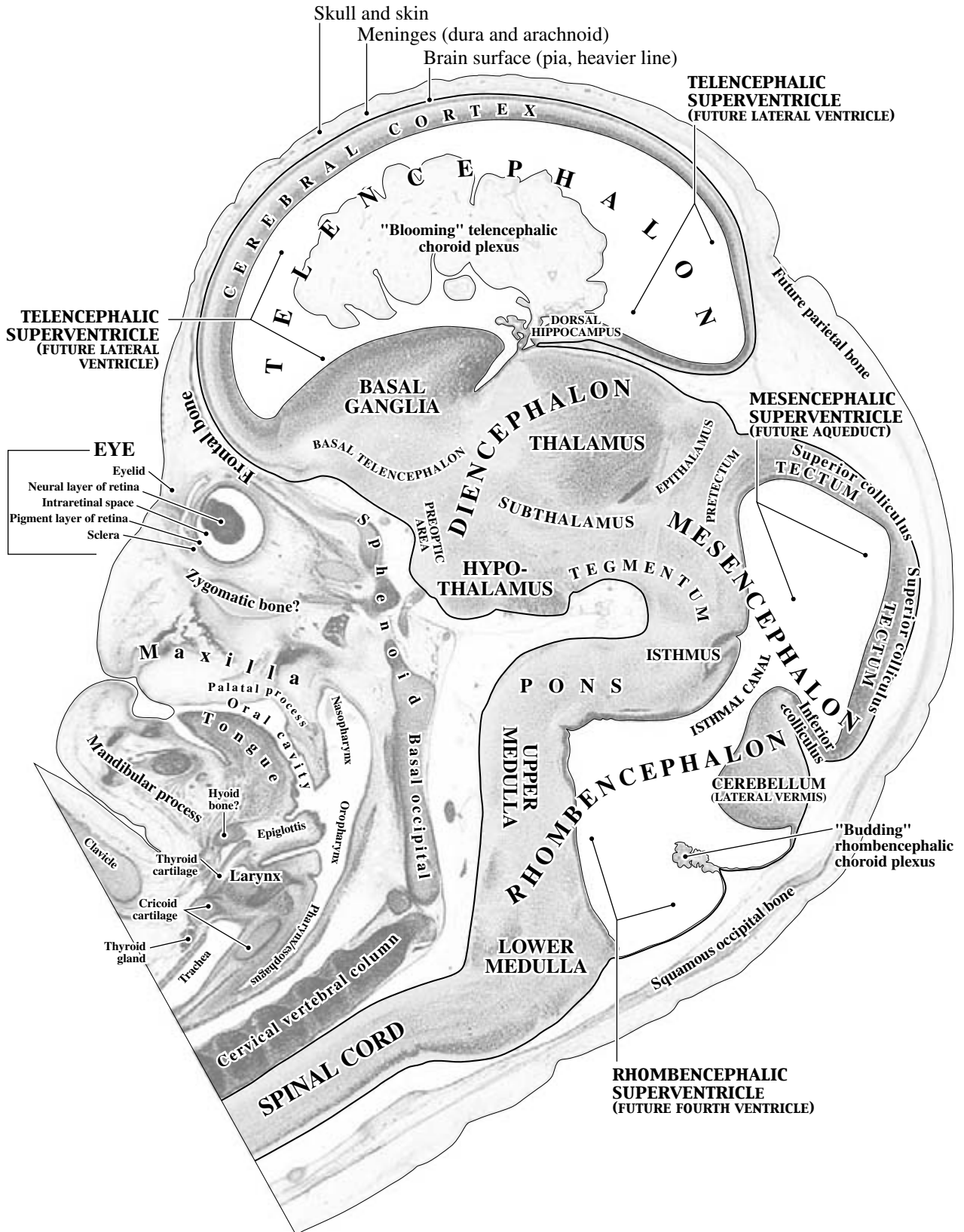
1. Medullary
2. Pontine
3. Mesencephalic
4. Diencephalic





Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 62B



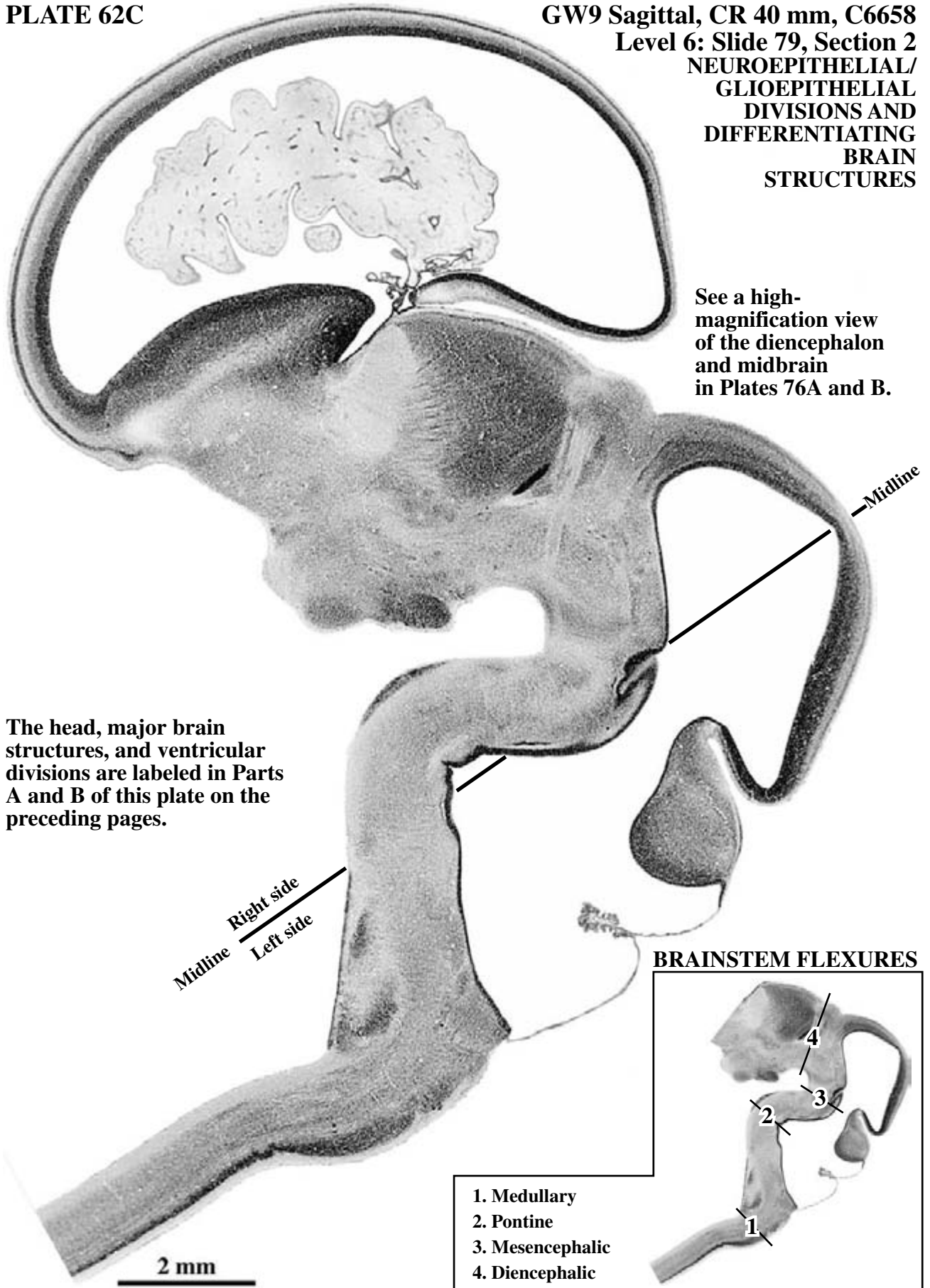
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

PLATE 62C

GW9 Sagittal, CR 40 mm, C6658
Level 6: Slide 79, Section 2
NEUROEPITHELIAL/
GLIOEPITHELIAL
DIVISIONS AND
DIFFERENTIATING
BRAIN
STRUCTURES

See a high-
magnification
view of the
diencephalon
and midbrain
in Plates 76A and B.

The head, major brain
structures, and ventricular
divisions are labeled in Parts
A and B of this plate on the
preceding pages.

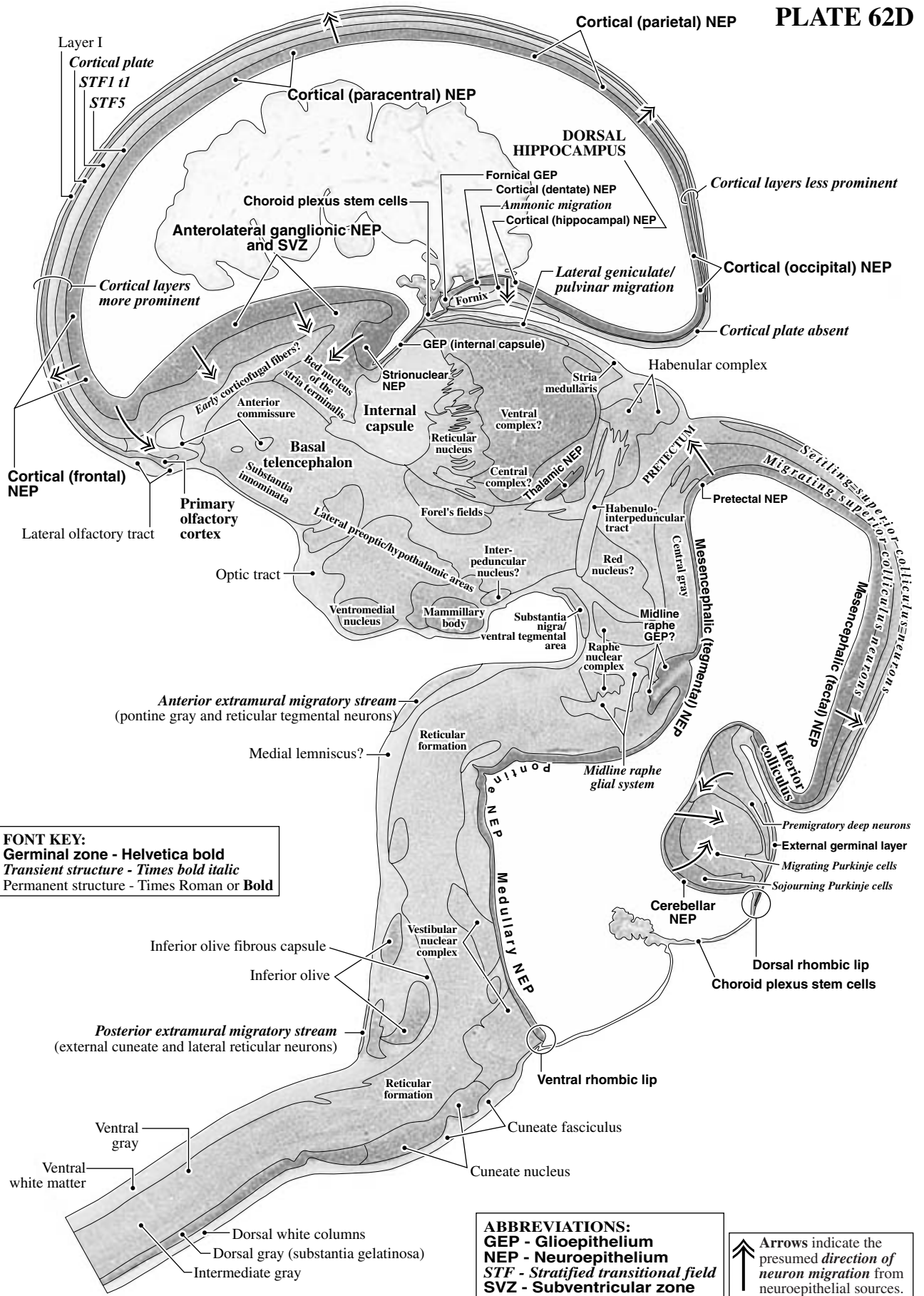


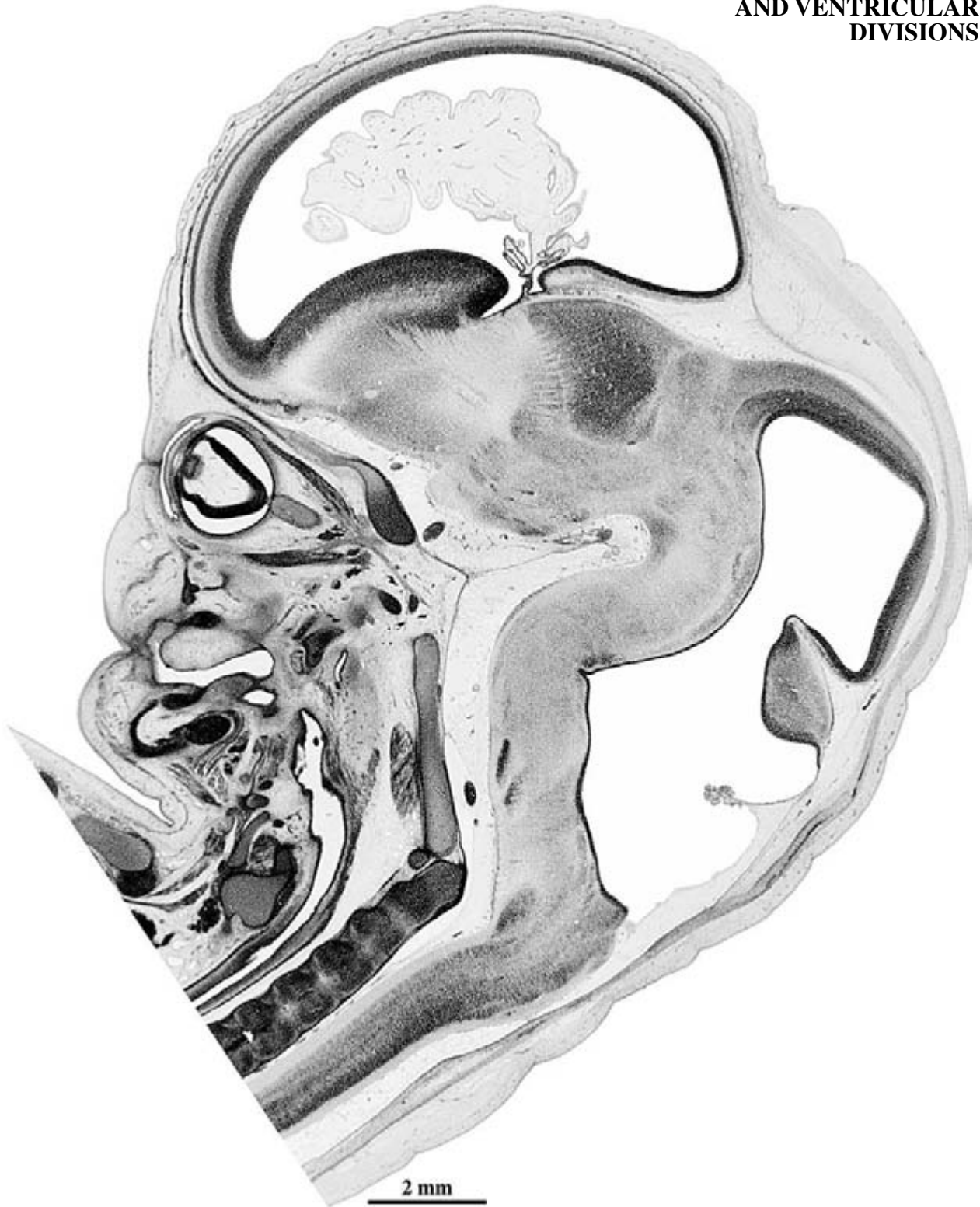
Right side
Left side
Midline

BRAINSTEM FLEXURES

- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

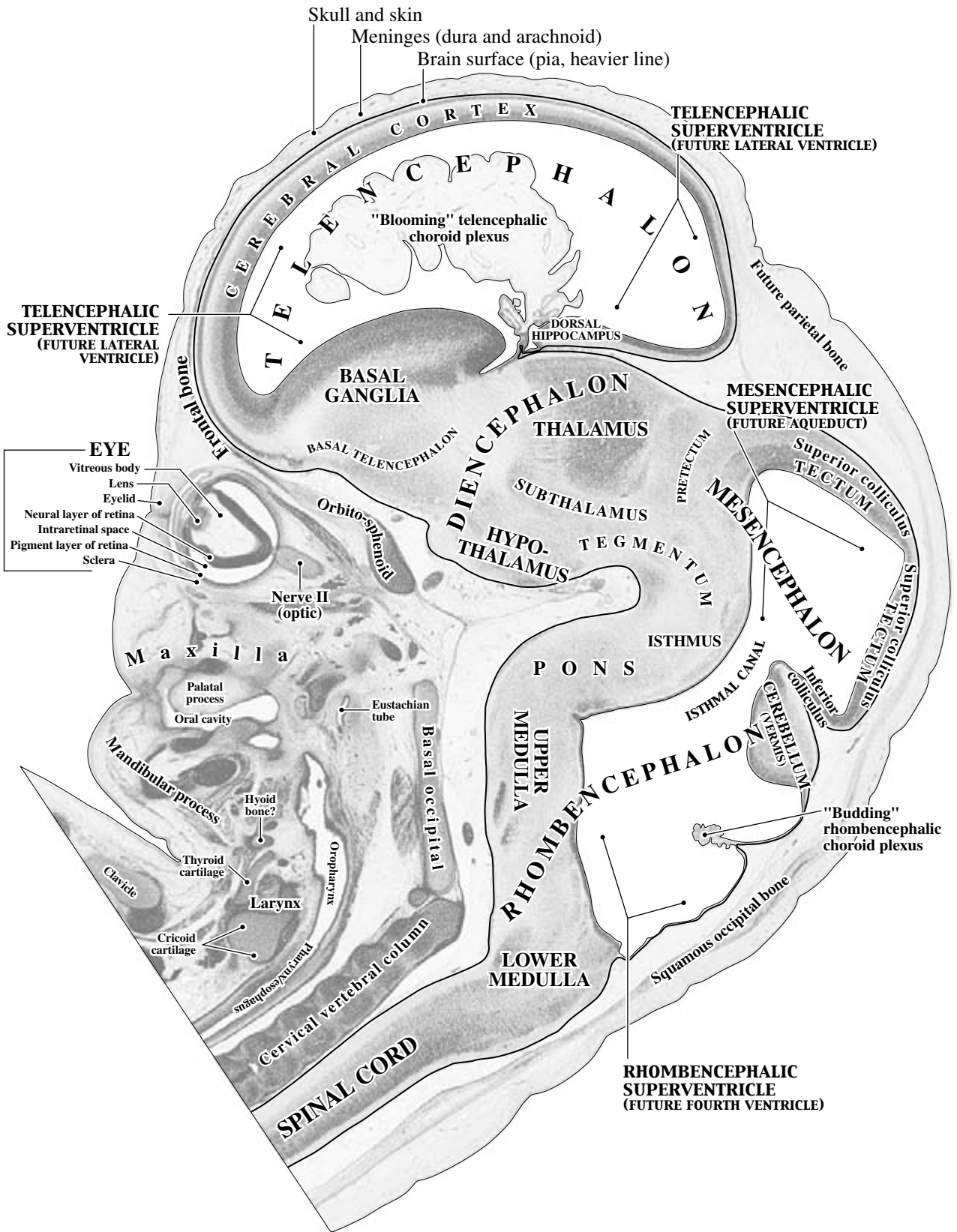
2 mm





Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 63B



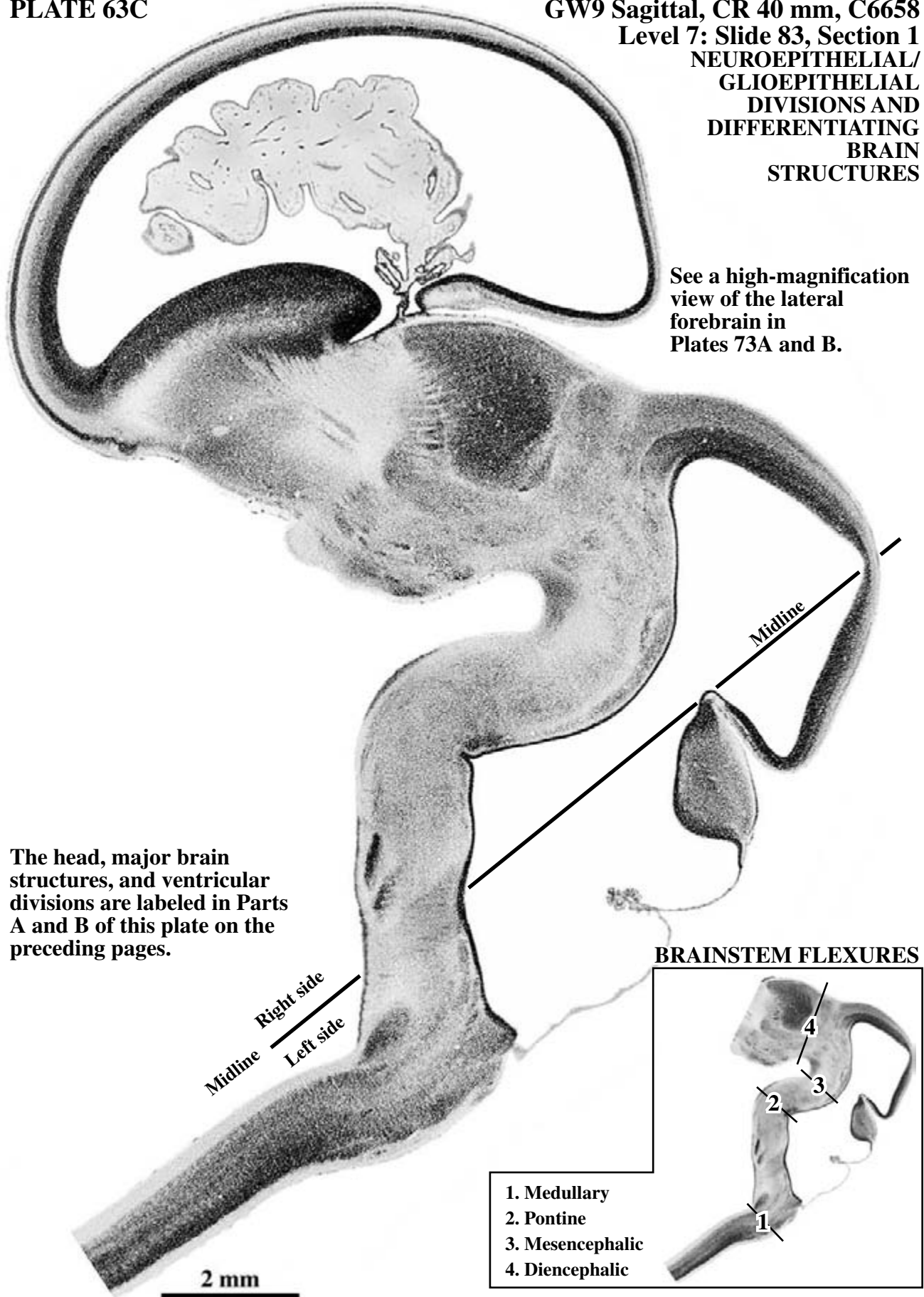
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

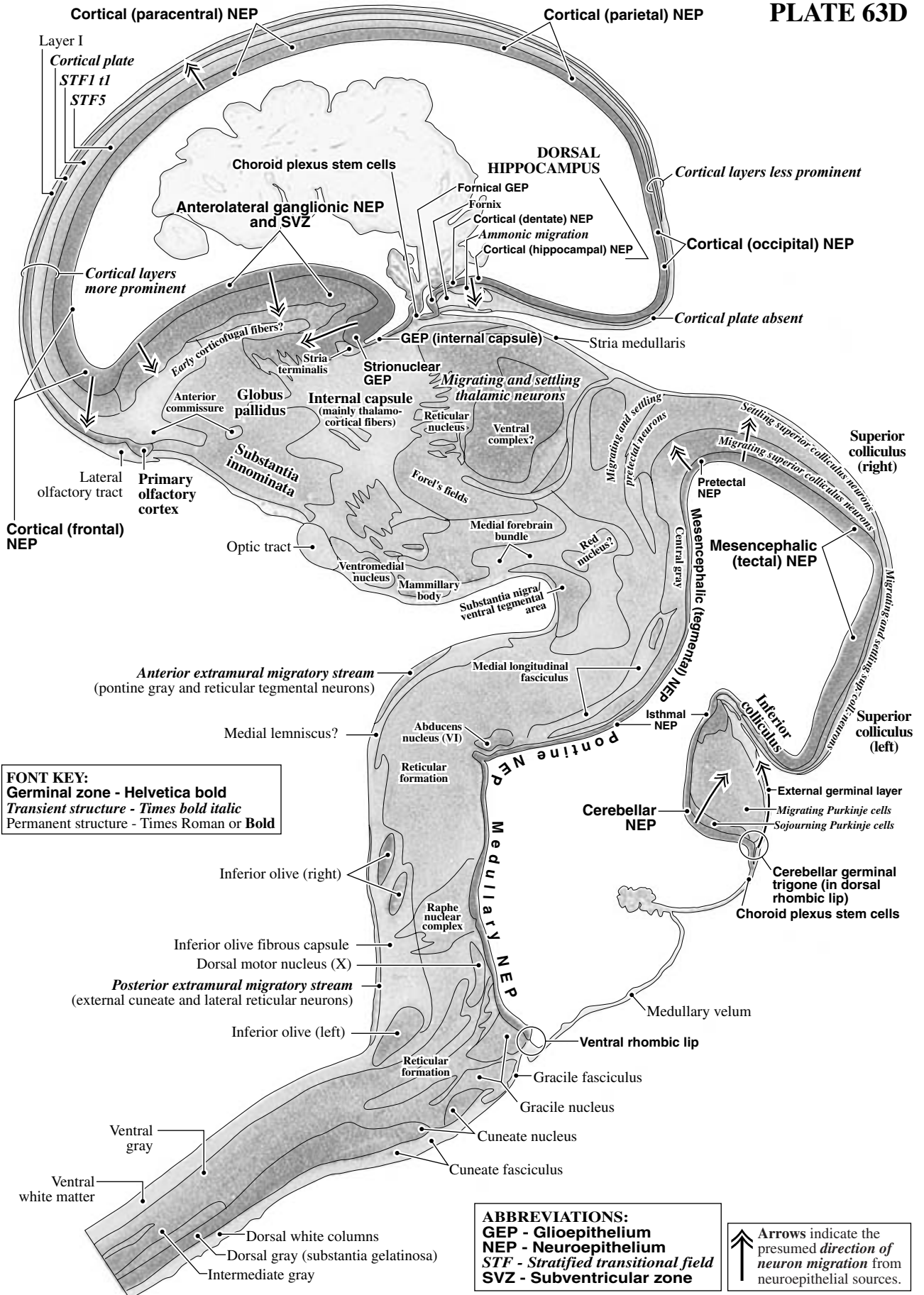
PLATE 63C

GW9 Sagittal, CR 40 mm, C6658
Level 7: Slide 83, Section 1
NEUROEPITHELIAL/
GLIOEPITHELIAL
DIVISIONS AND
DIFFERENTIATING
BRAIN
STRUCTURES

See a high-magnification
view of the lateral
forebrain in
Plates 73A and B.

The head, major brain
structures, and ventricular
divisions are labeled in Parts
A and B of this plate on the
preceding pages.





FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

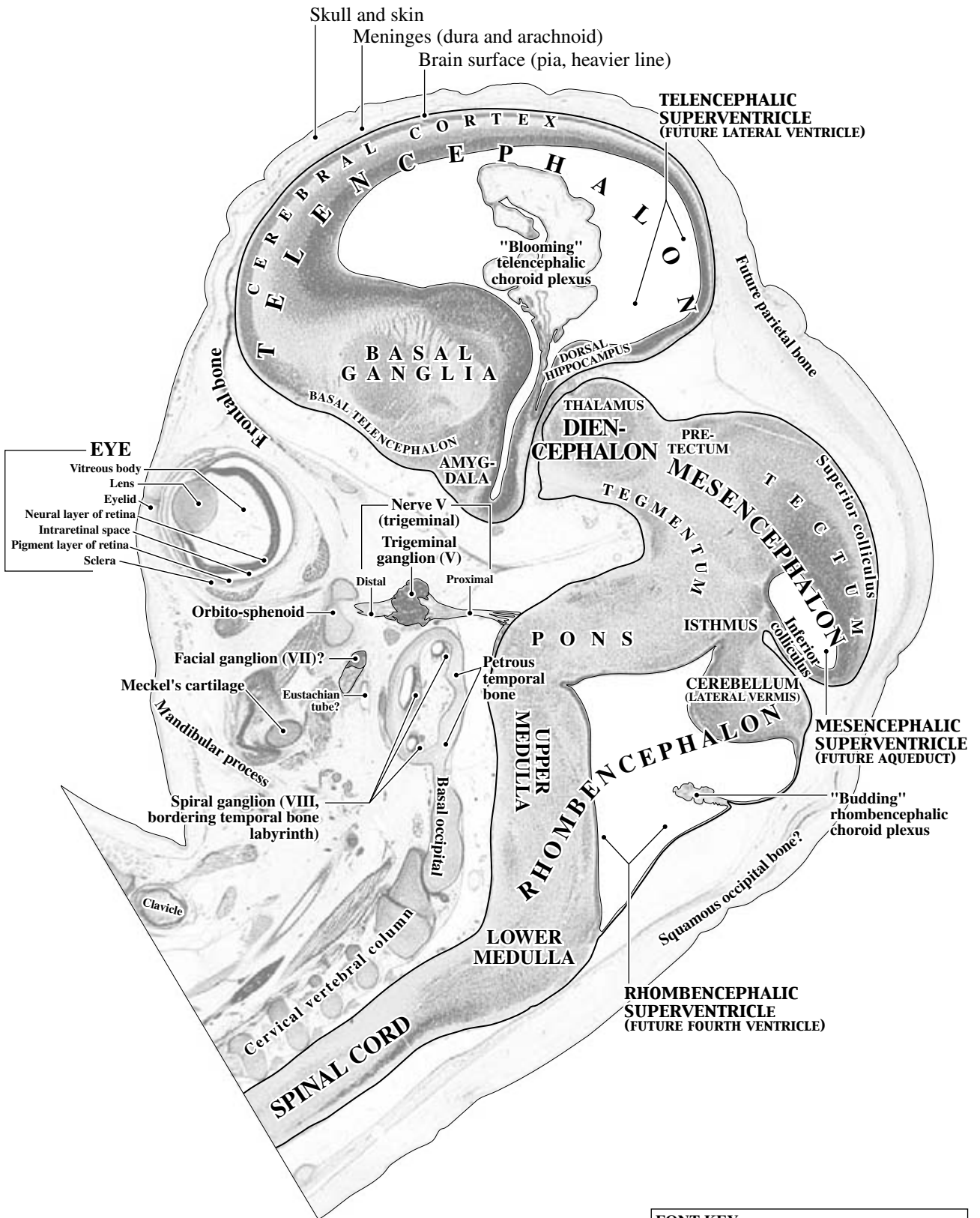
ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.



Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 64B



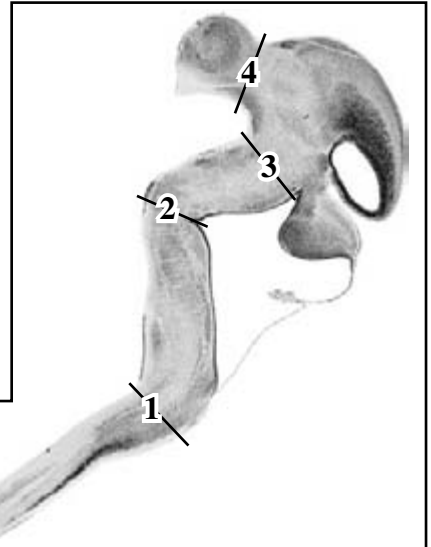
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **B**old CAPITALS
 All other structures - Times Roman or **B**old

GW9 Sagittal, CR 40 mm, C6658
 Level 8: Slide 95, Section 1
 NEUROEPITHELIAL/
 GLIOEPITHELIAL
 DIVISIONS AND
 DIFFERENTIATING
 BRAIN STRUCTURES

See a high-magnification
 view of the hippocampus
 and thalamus
 in Plates 72A and B.

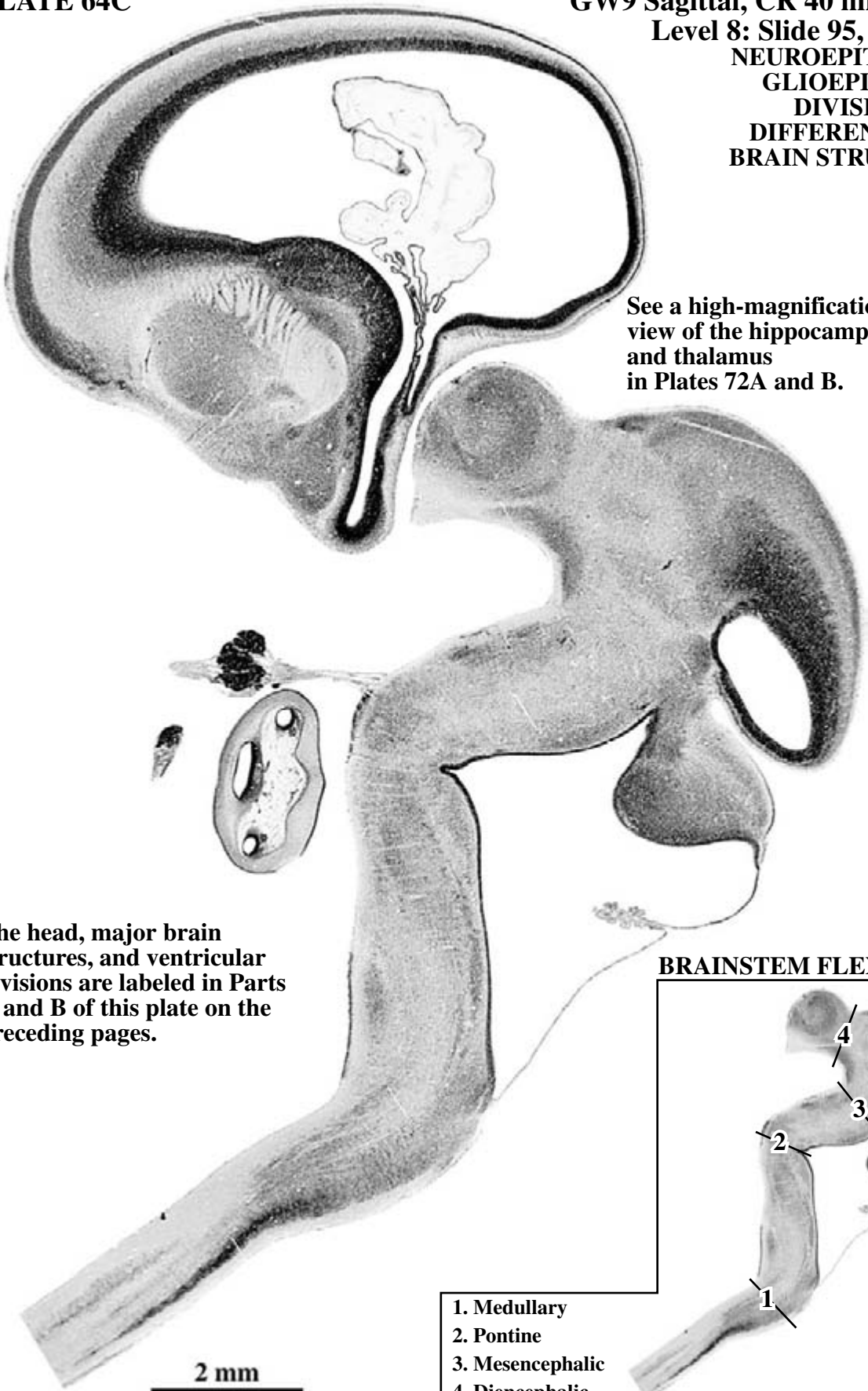
The head, major brain
 structures, and ventricular
 divisions are labeled in Parts
 A and B of this plate on the
 preceding pages.

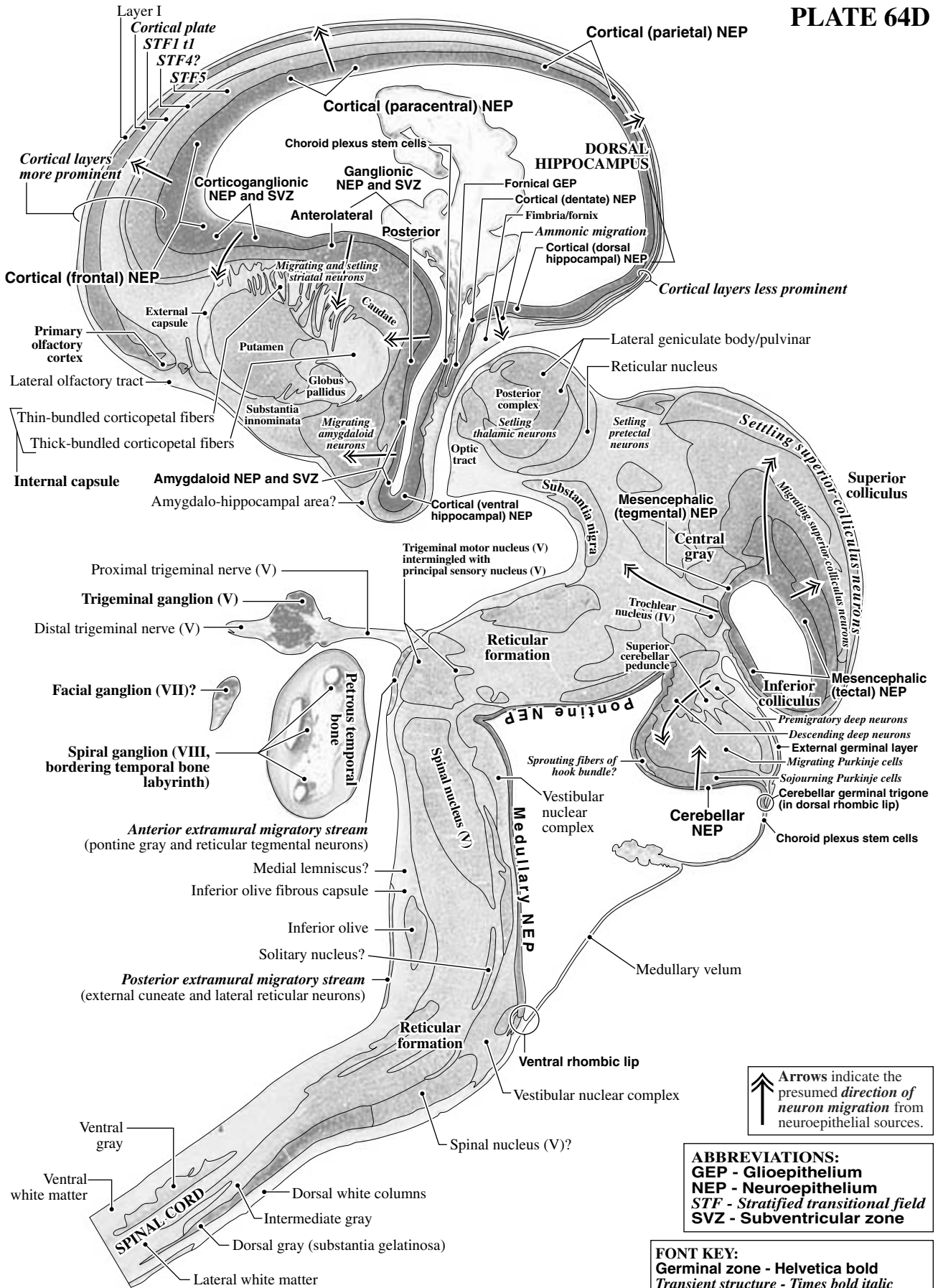
BRAINSTEM FLEXURES



- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

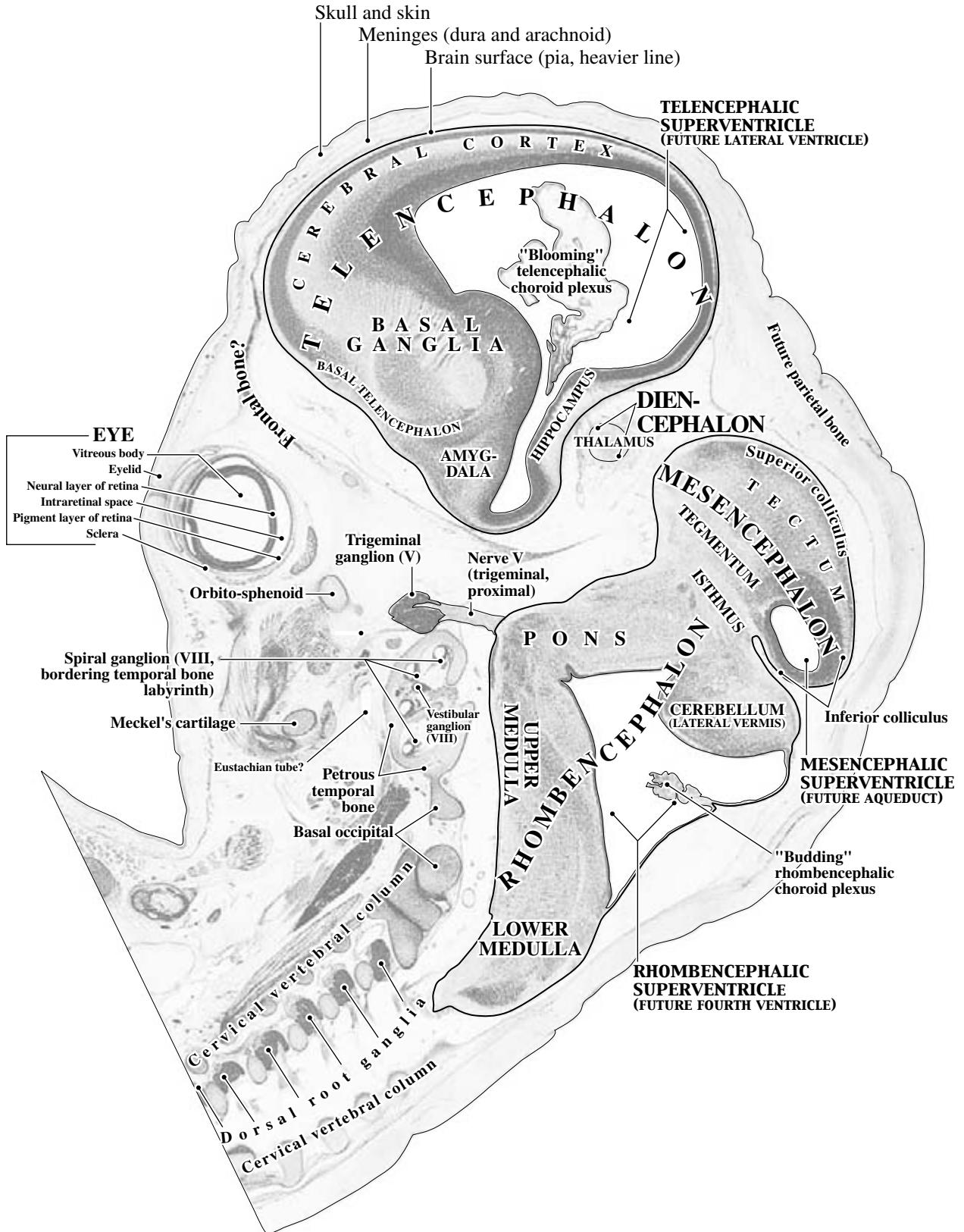
2 mm



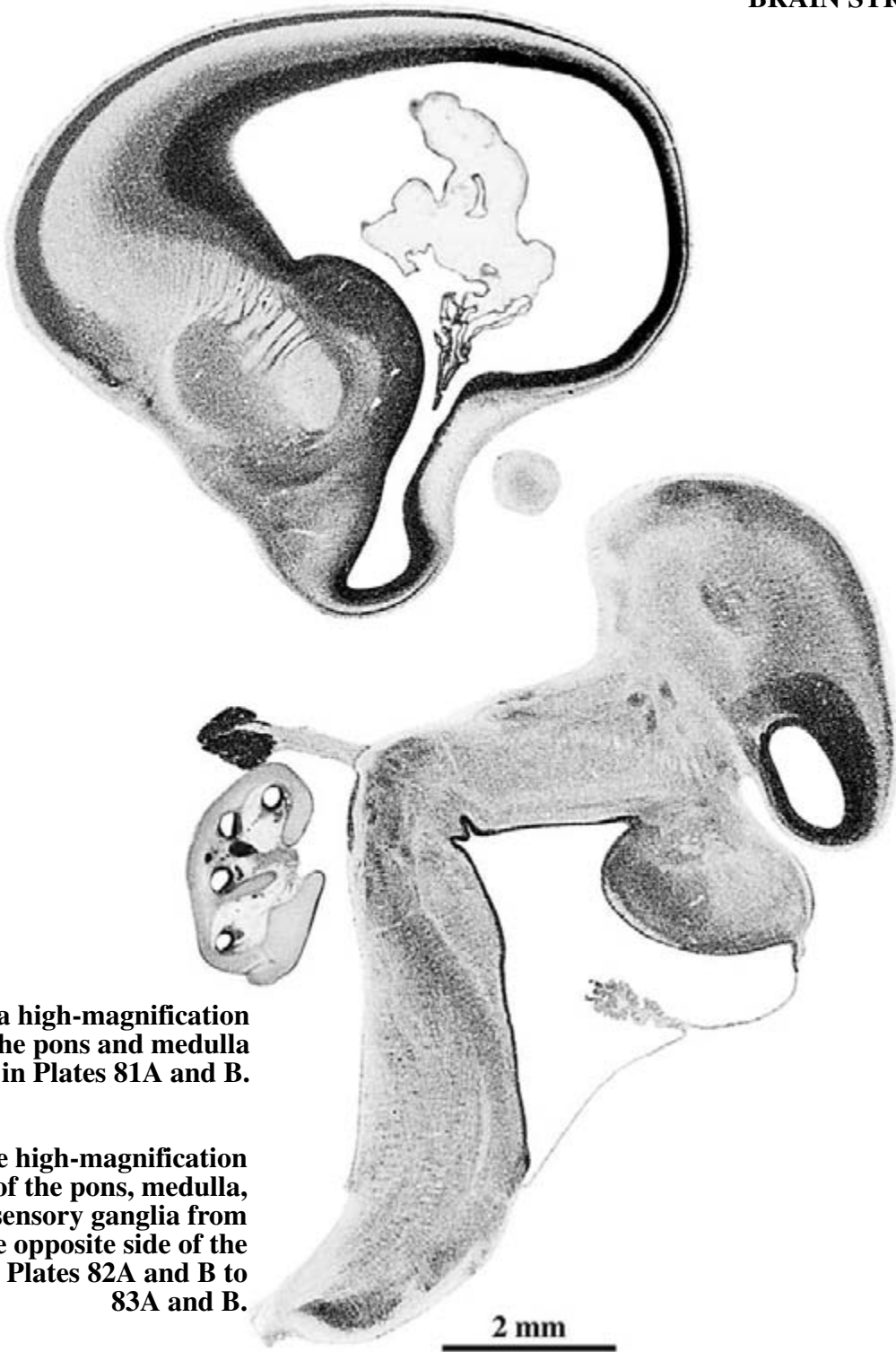




Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



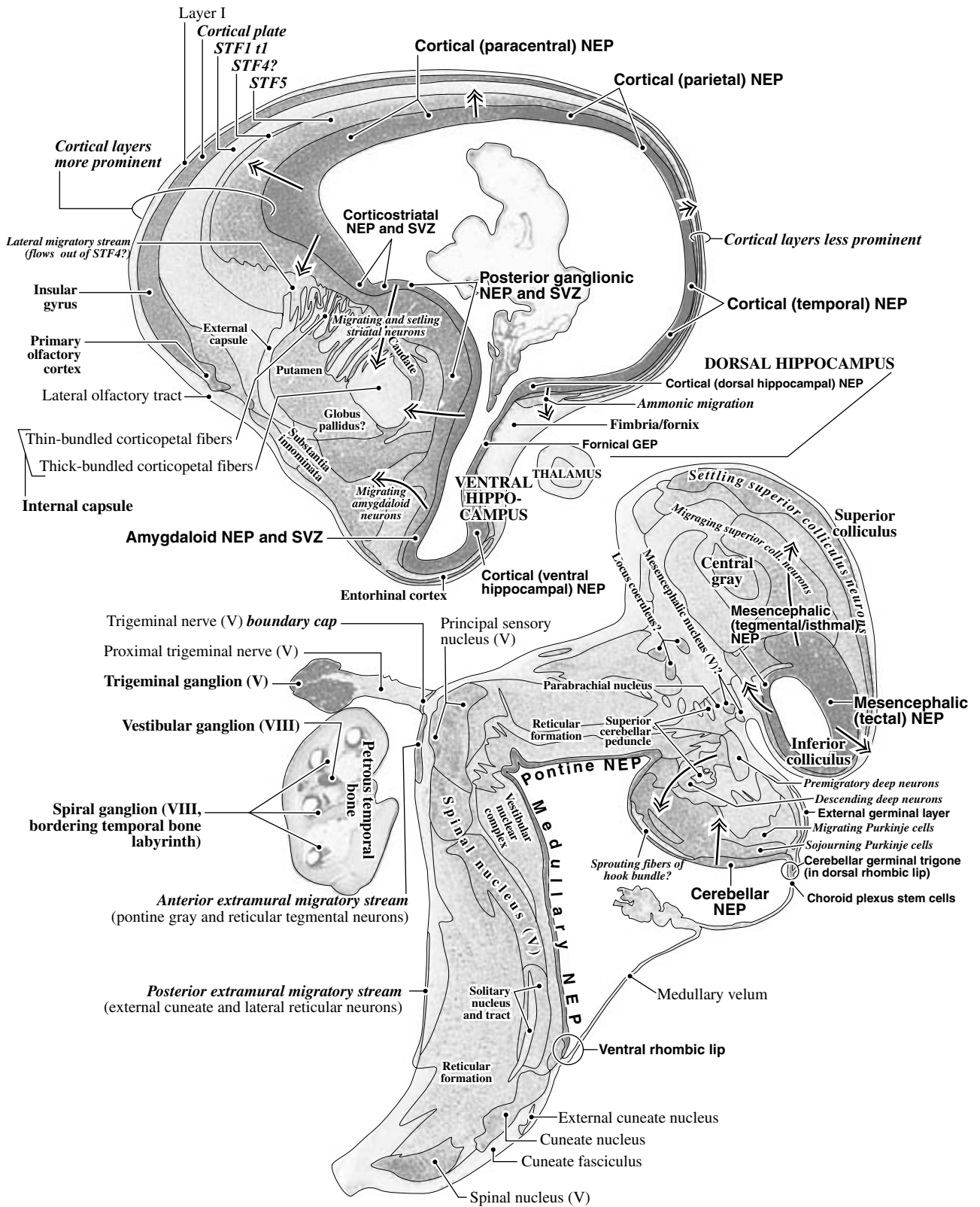
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See a high-magnification view of the pons and medulla in Plates 81A and B.

See high-magnification views of the pons, medulla, and sensory ganglia from the opposite side of the brain in Plates 82A and B to 83A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

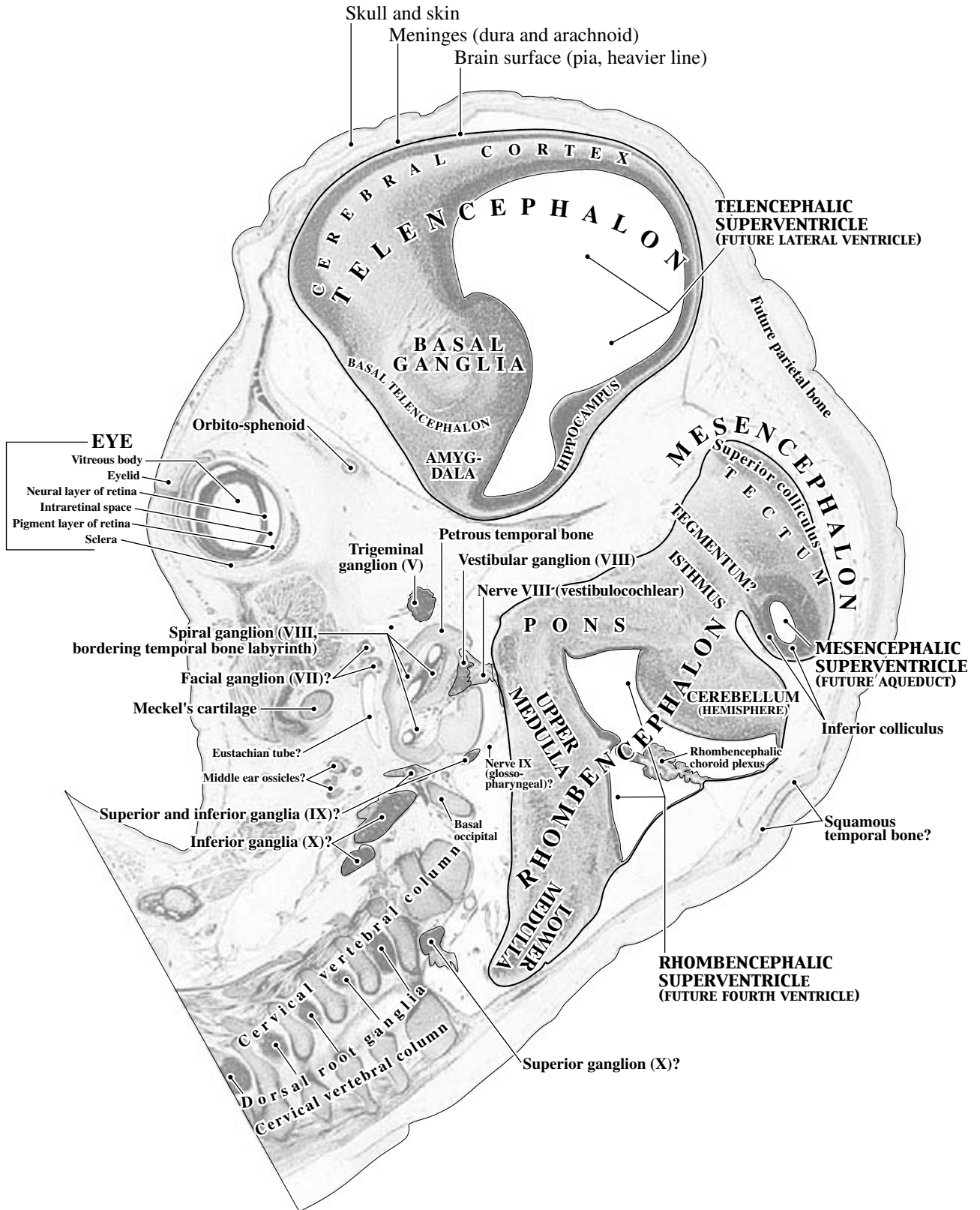
ABBREVIATIONS:
GEP - Glioeptelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

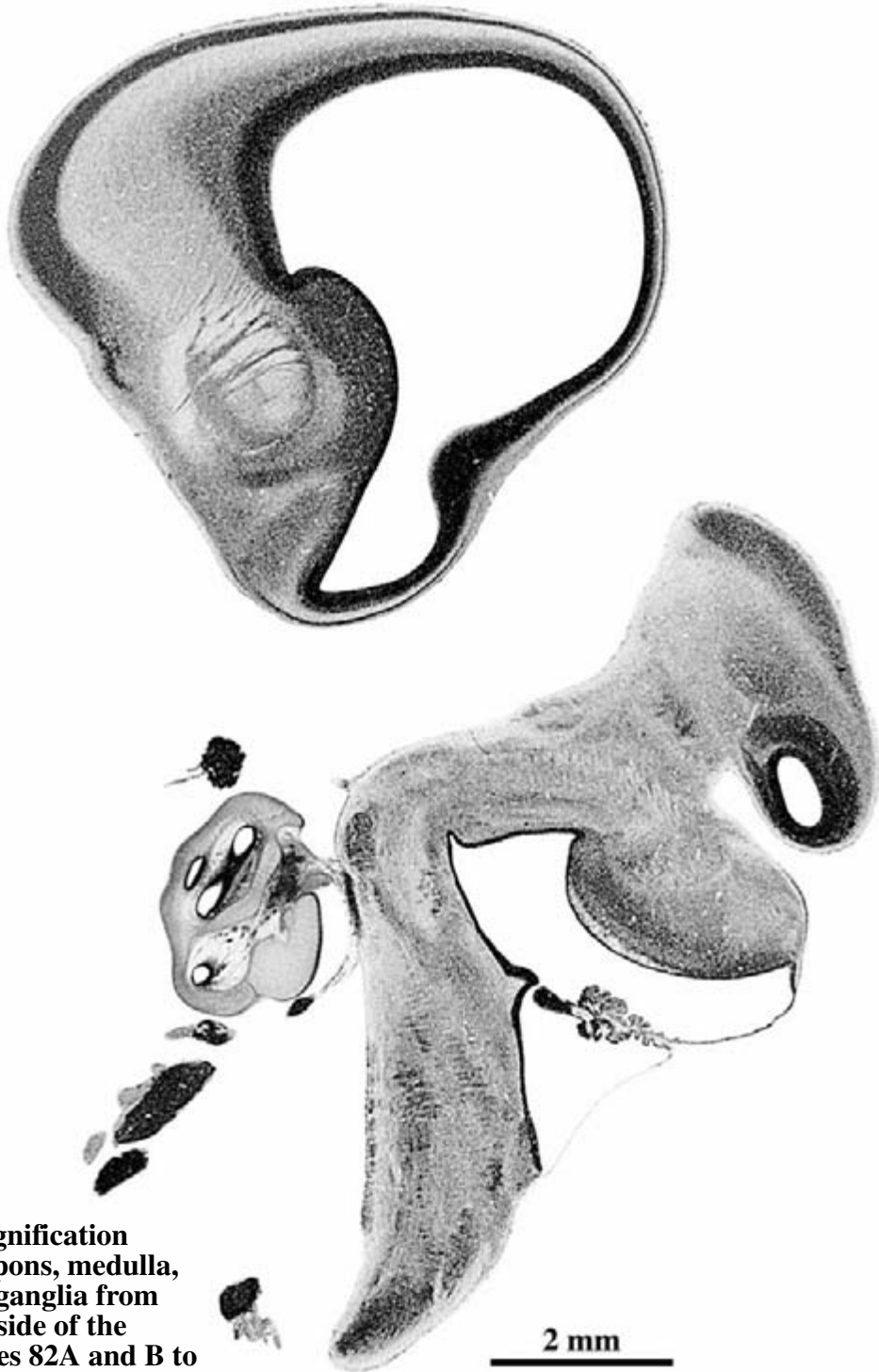


Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 66B

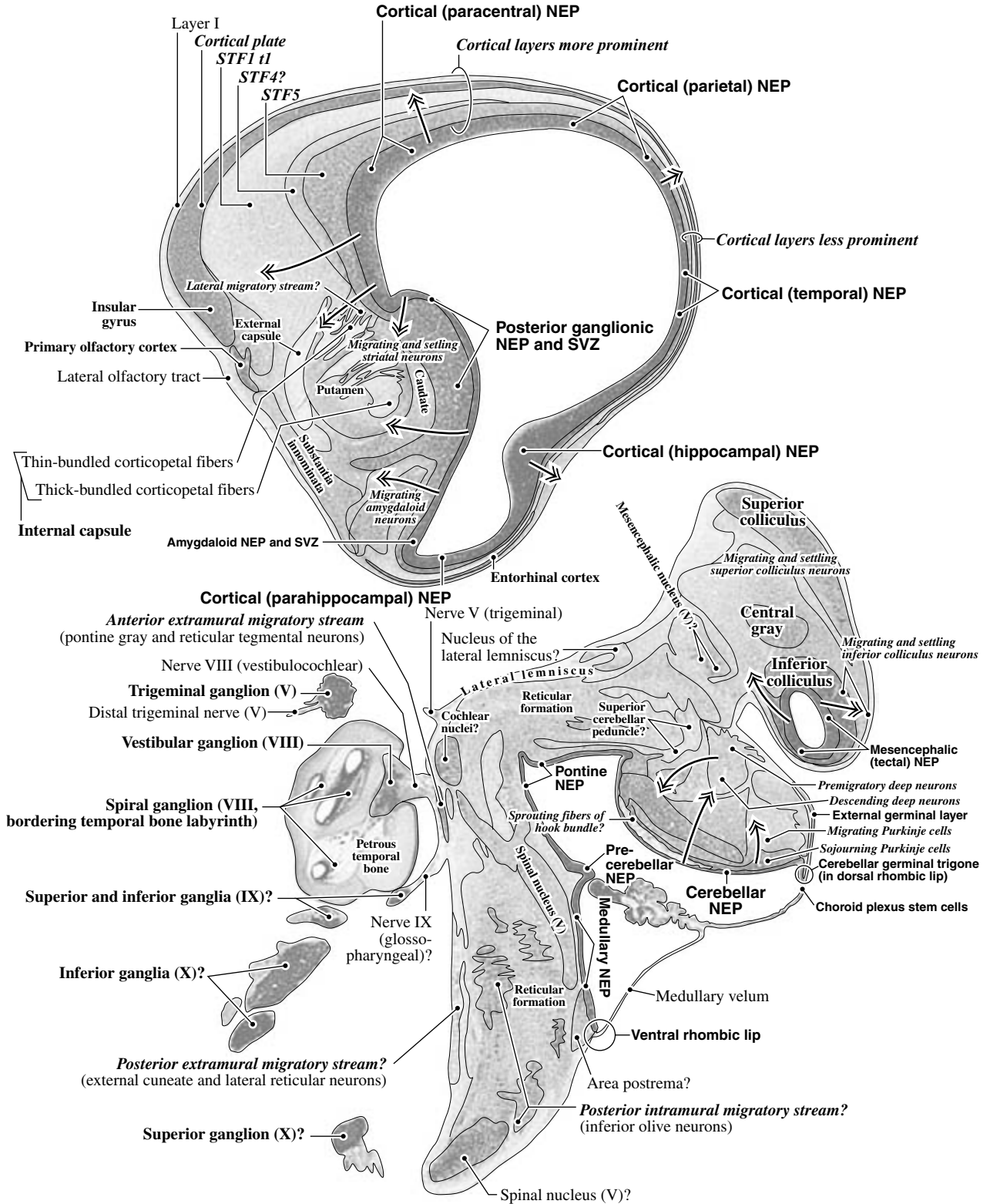


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See high-magnification views of the pons, medulla, and sensory ganglia from the opposite side of the brain in Plates 82A and B to 83A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

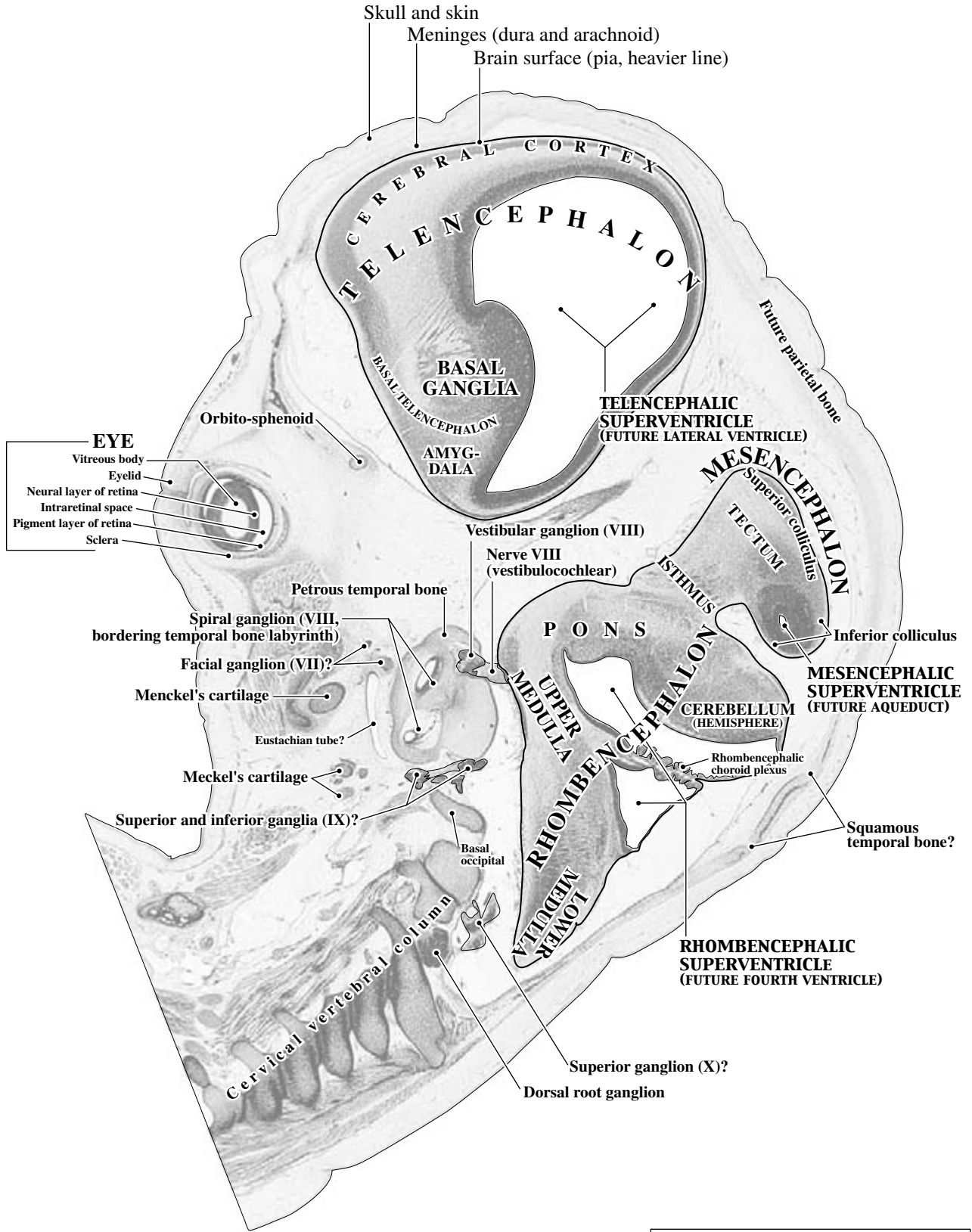
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

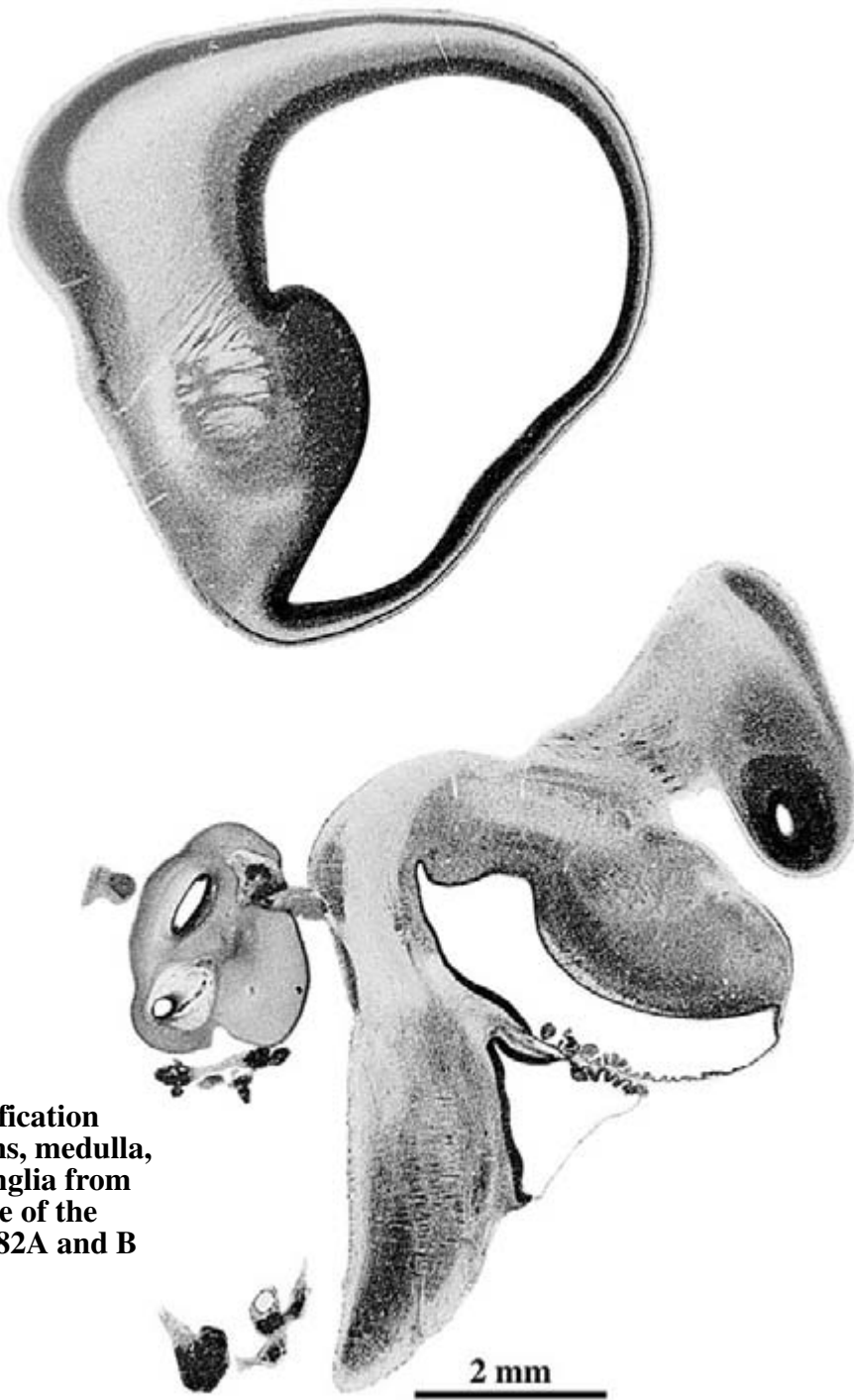


Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

PLATE 67B

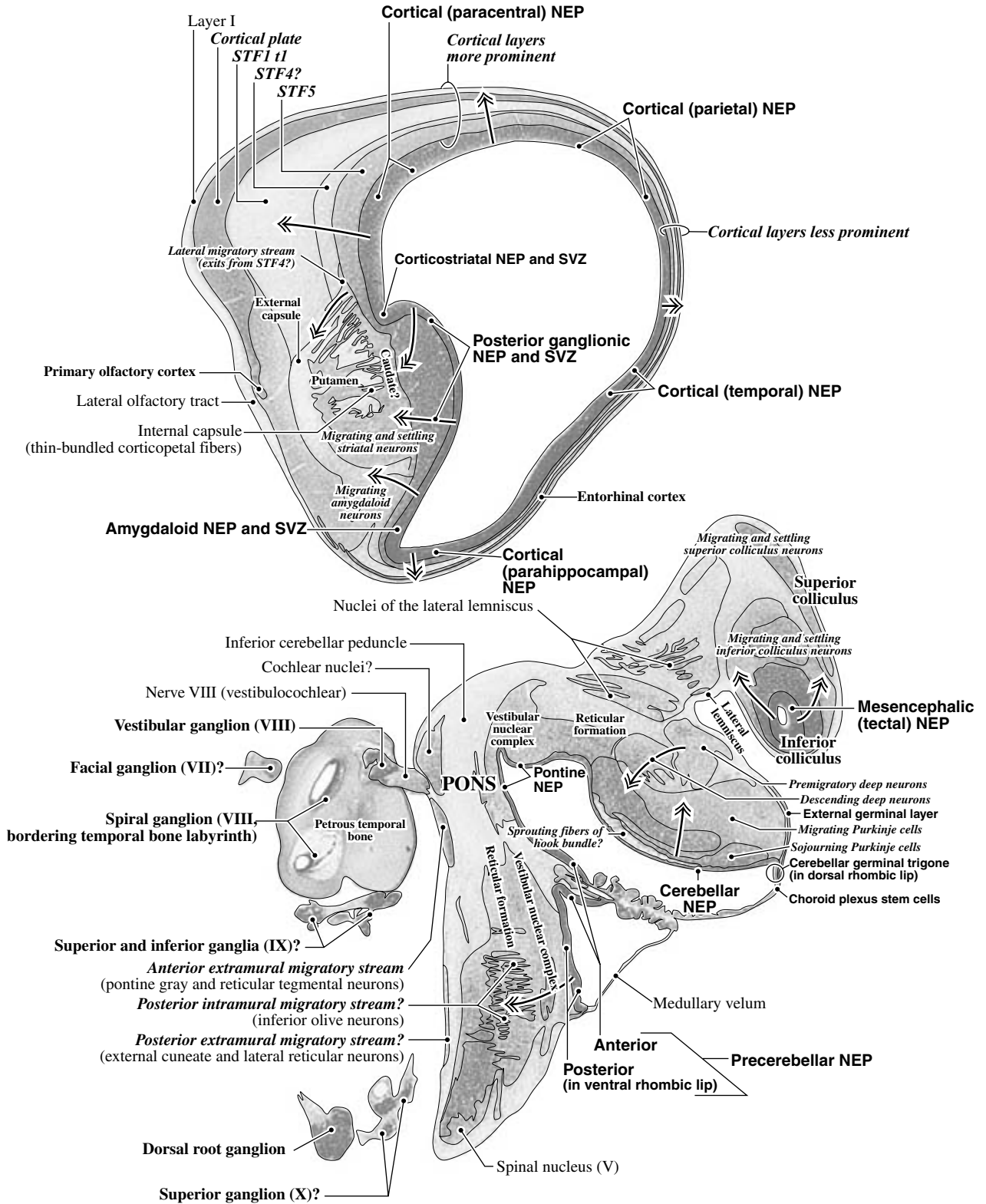


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See high-magnification views of the pons, medulla, and sensory ganglia from the opposite side of the brain in Plates 82A and B to 83A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

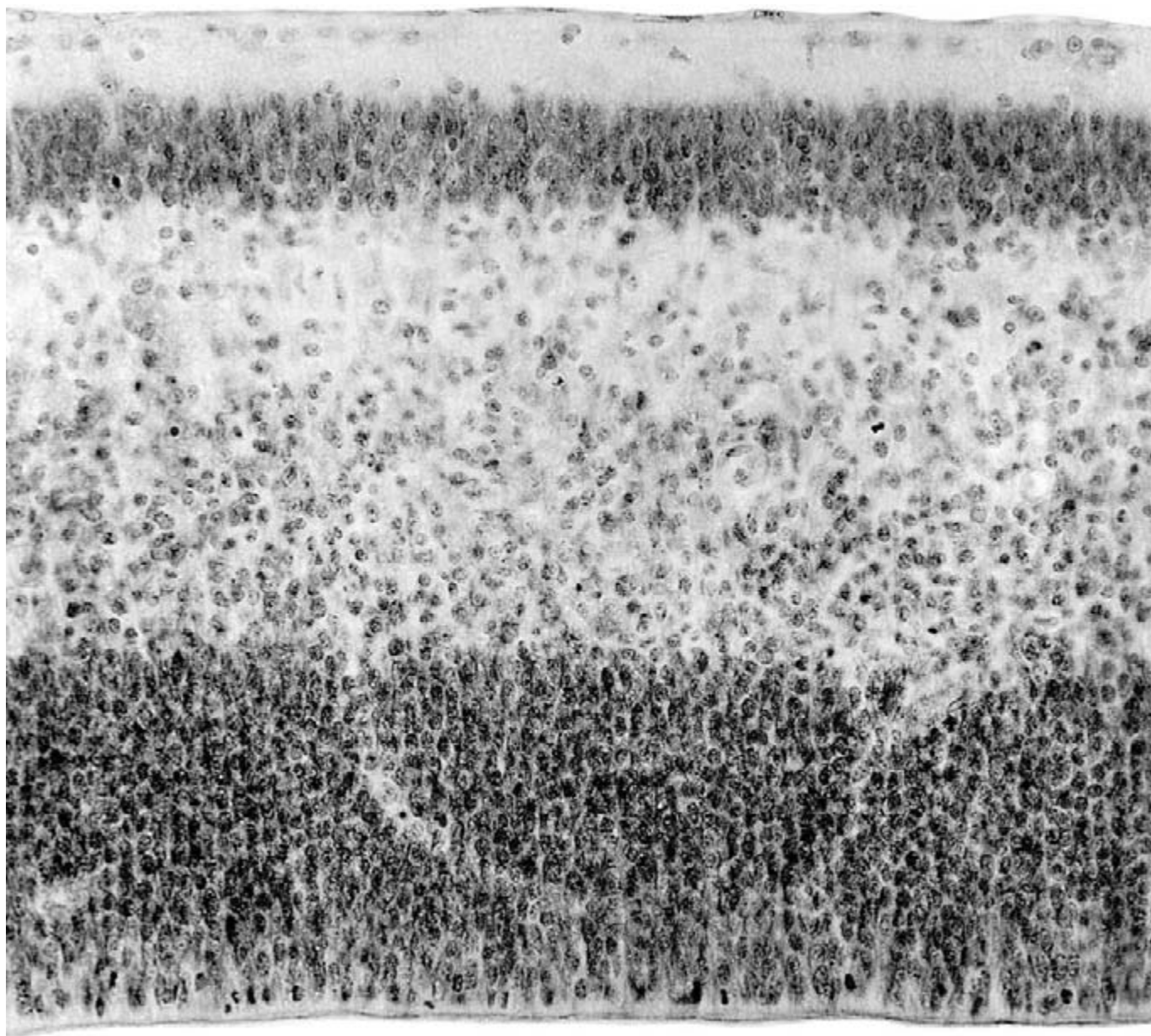
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 68A

**GW9 Sagittal
CR 40 mm, C6658
Level 4:
Slide 71, Section 2**

**DORSAL
CORTEX**



See the entire section in Plates 60A-D.

0.05 mm

PLATE 68B

Layer I contains Cajal-Retzius cells settled near the pia.

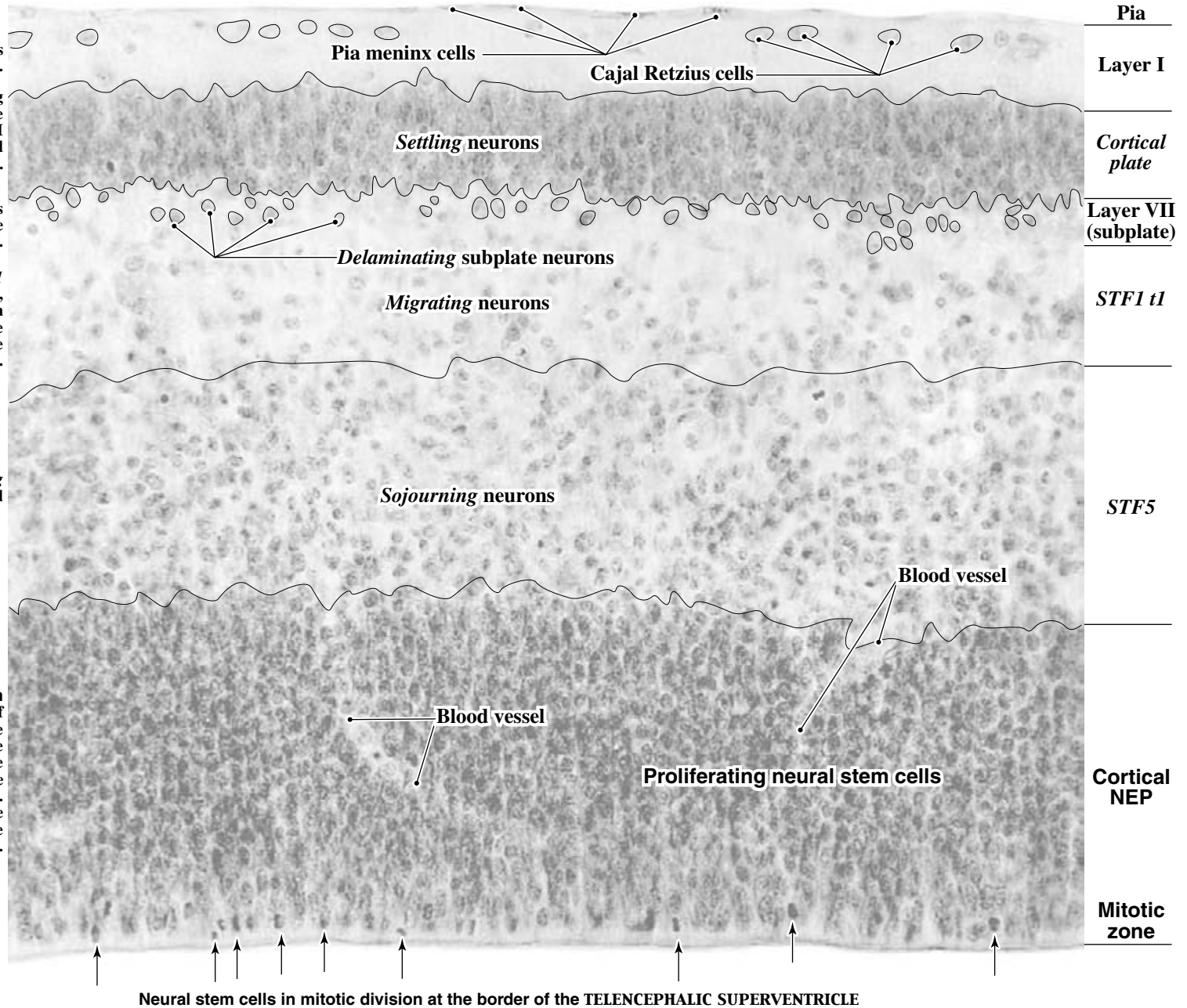
The cortical plate contains settling neurons, mainly those in future layer VI and some in layer VII that have not yet migrated downward into the subplate.

The subplate contains neurons that originally resided in the cortical plate.

Stratified transitional field (STF) 1 contains migrating neurons, mainly those that will settle in cortical layer VI. STF1 is in the early t1 stage when cells are plentiful among the fibers.

STF 5 contains sojourning neurons, mainly those that will settle in cortical layers V and VI.

The cortical neuroepithelium (NEP) is a pseudostratified layer of neural stem cells. Arrows indicate mitotic figures. The majority are at the ventricular surface, where nuclei migrate to divide within the tall and thin columnar cytoplasm. At this stage of development, large numbers of cortical neurons are being generated.

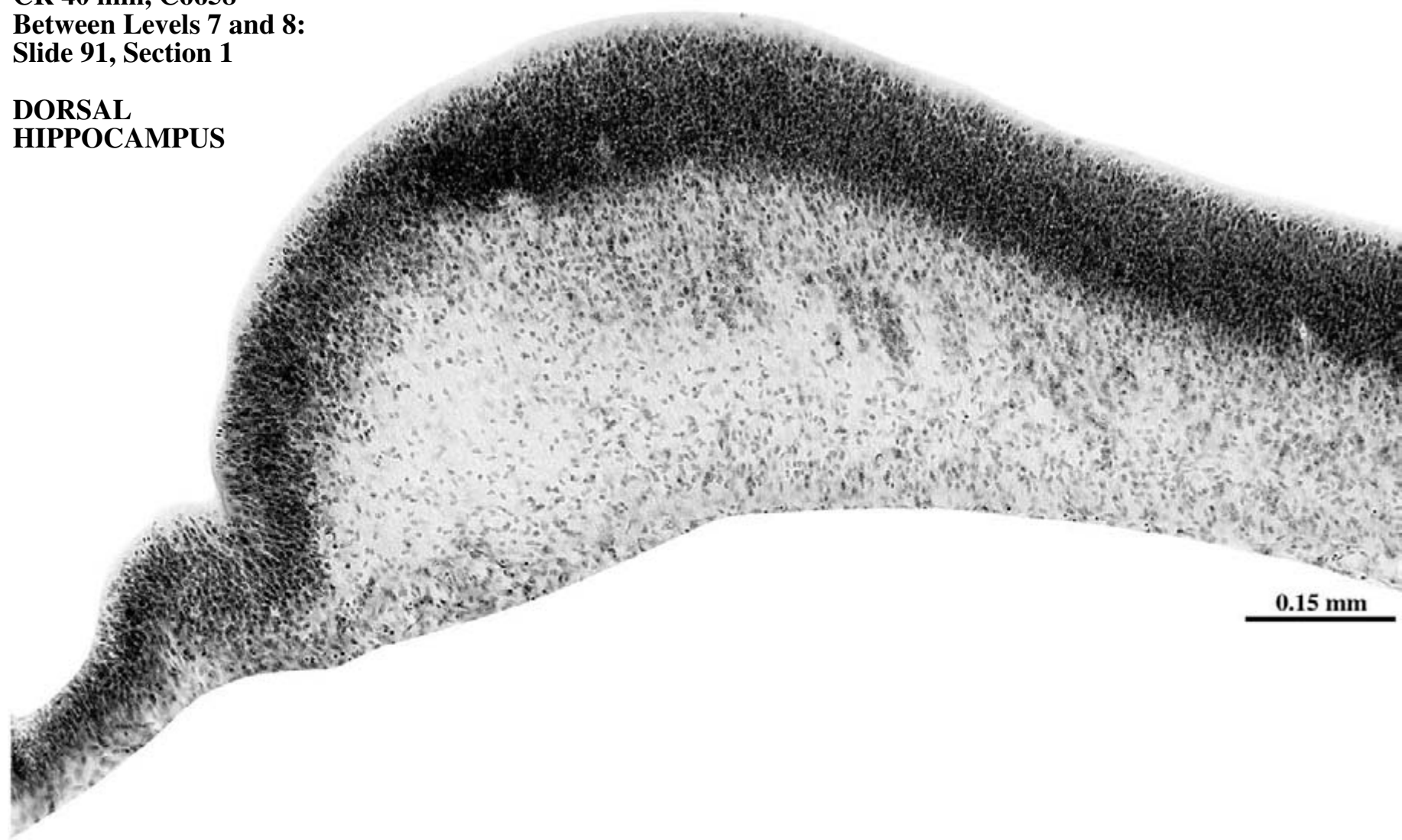


Neural stem cells in mitotic division at the border of the TELECEPHALIC SUPERVENTRICLE

PLATE 69A

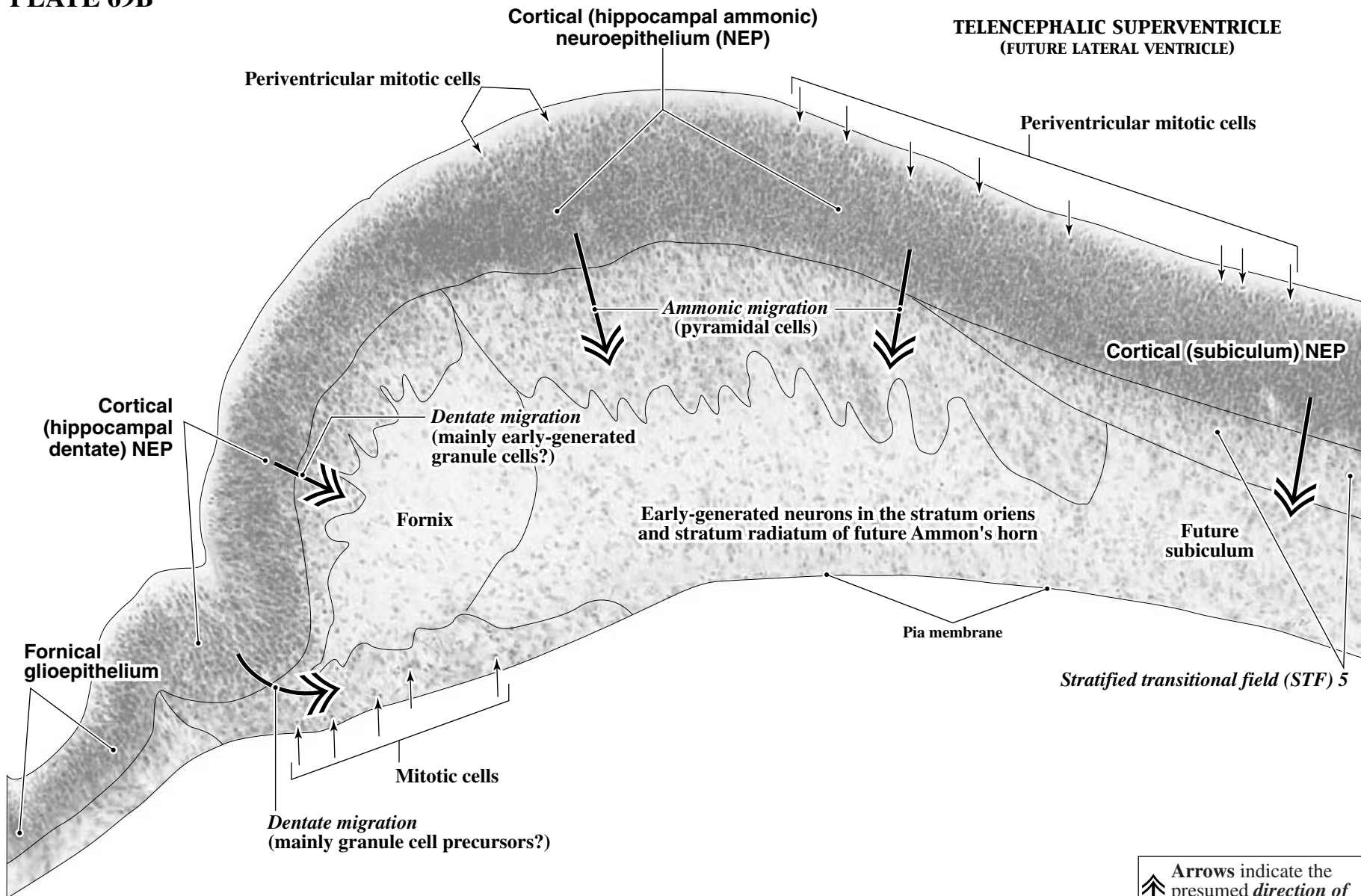
**GW9 Sagittal
CR 40 mm, C6658
Between Levels 7 and 8:
Slide 91, Section 1**

**DORSAL
HIPPOCAMPUS**



See a low-magnification view of Level 7 in Plates 63A-D, Level 8 in Plates 64A-D.

PLATE 69B



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 70A

**GW9 Sagittal
CR 40 mm, C6658
Level 3:
Slide 67, Section 1**

**OLFACTORY BULB
AND SEPTUM**



**See the entire section
in Plates 59A-D.**

PLATE 70B

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
STF - Stratified transitional field

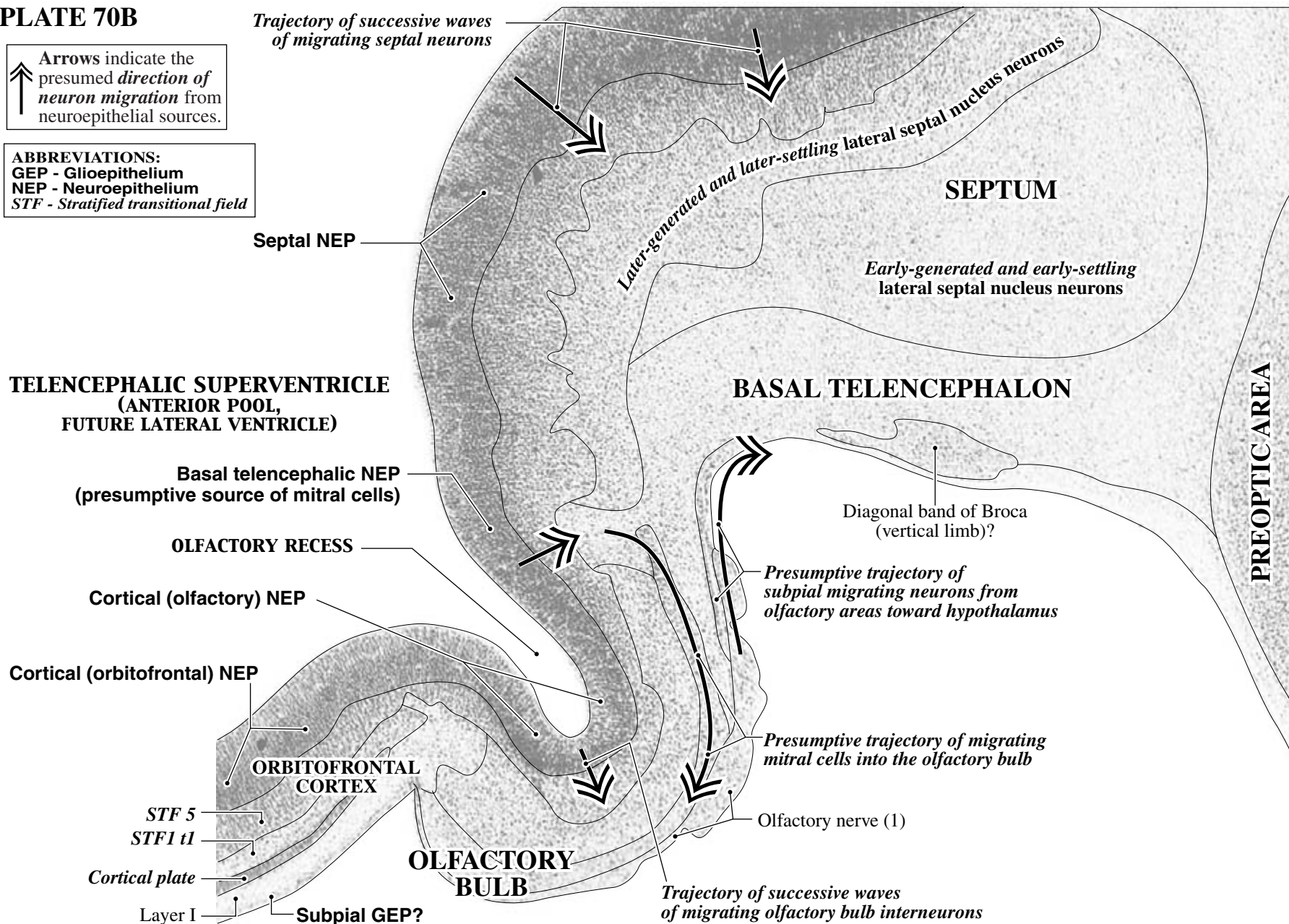
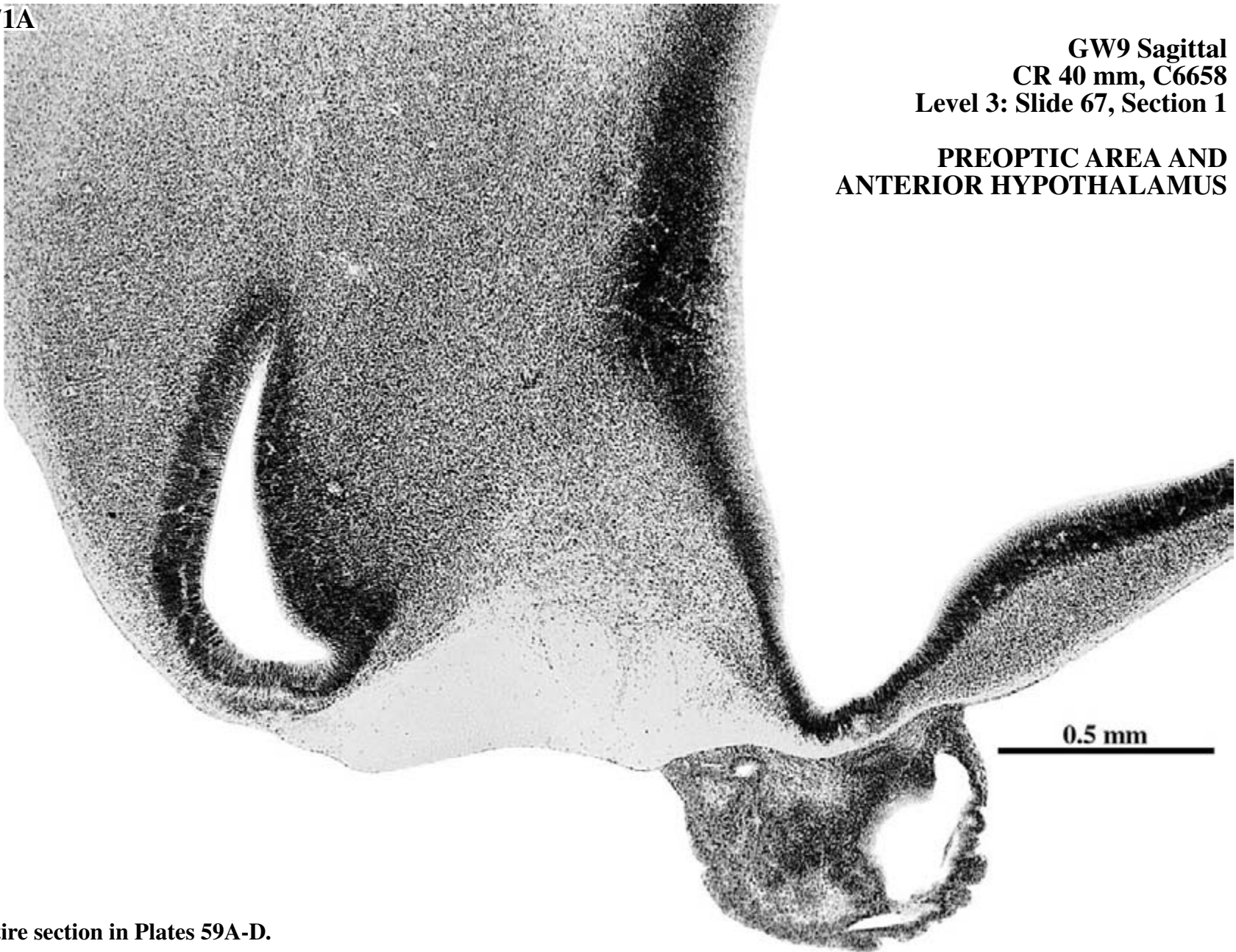


PLATE 71A

**GW9 Sagittal
CR 40 mm, C6658
Level 3: Slide 67, Section 1**

**PREOPTIC AREA AND
ANTERIOR HYPOTHALAMUS**



0.5 mm

See the entire section in Plates 59A-D.

PLATE 71B

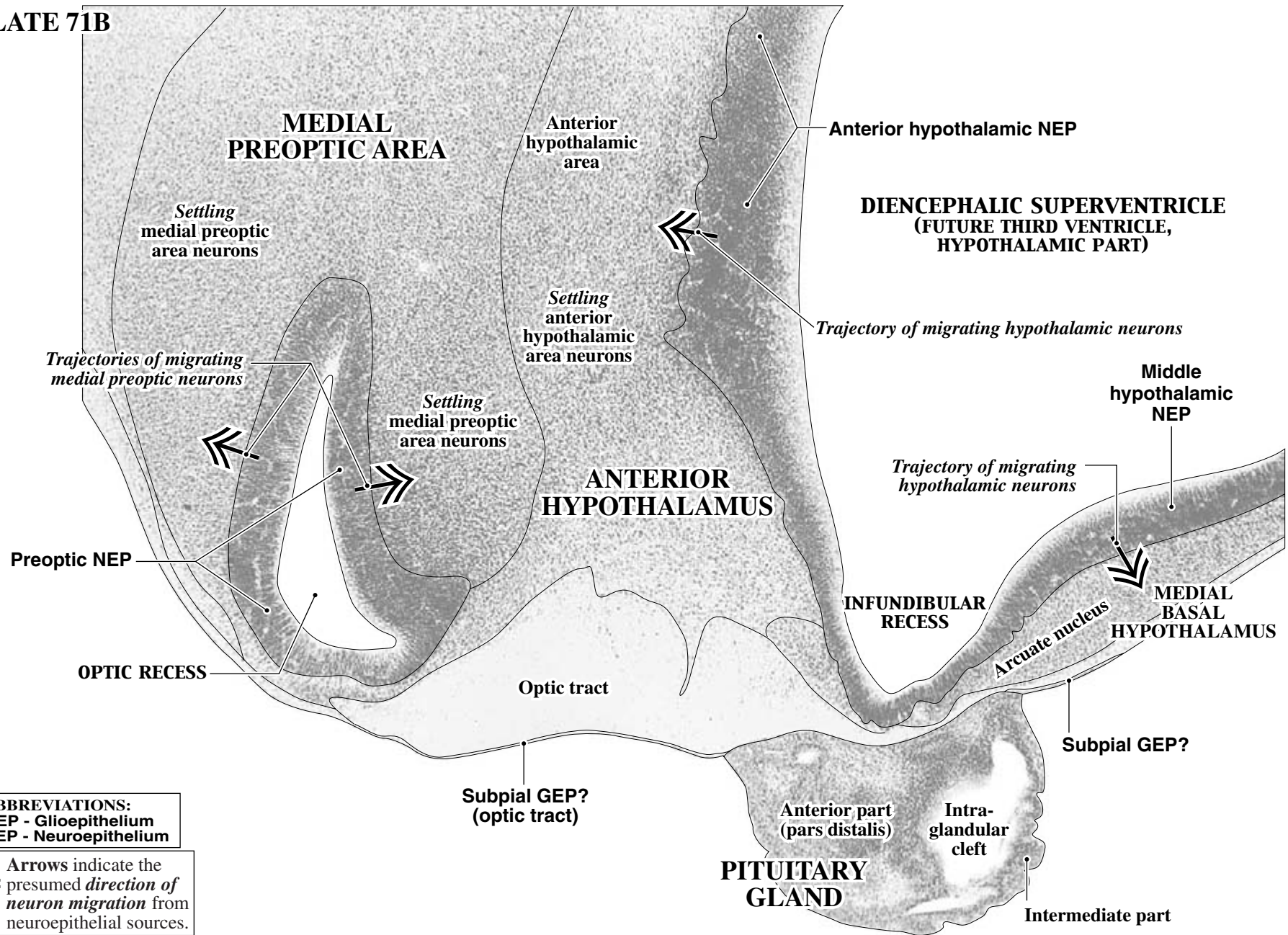
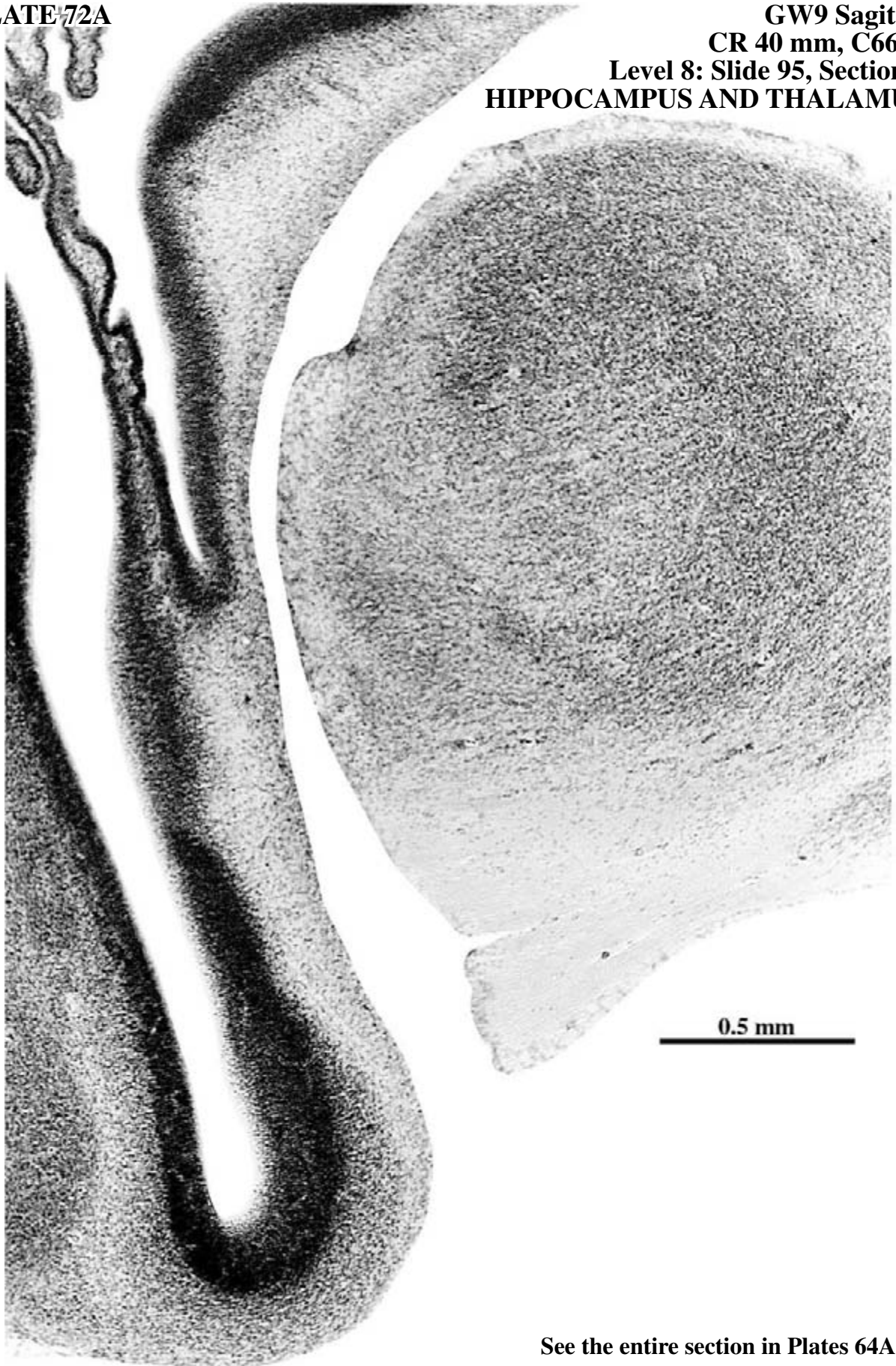


PLATE 72A

**GW9 Sagittal
CR 40 mm, C6658
Level 8: Slide 95, Section 1
HIPPOCAMPUS AND THALAMUS**



See the entire section in Plates 64A-D.

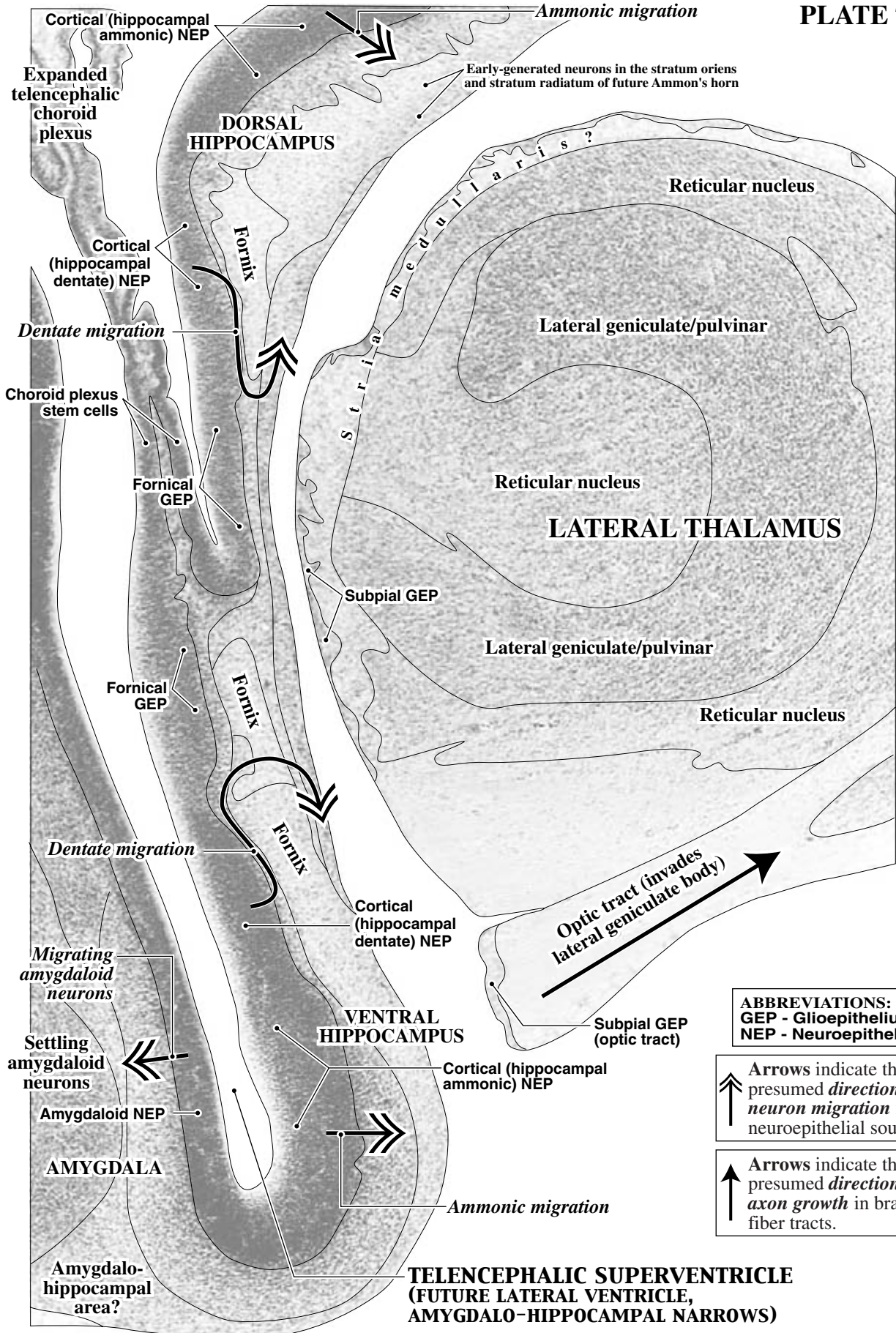
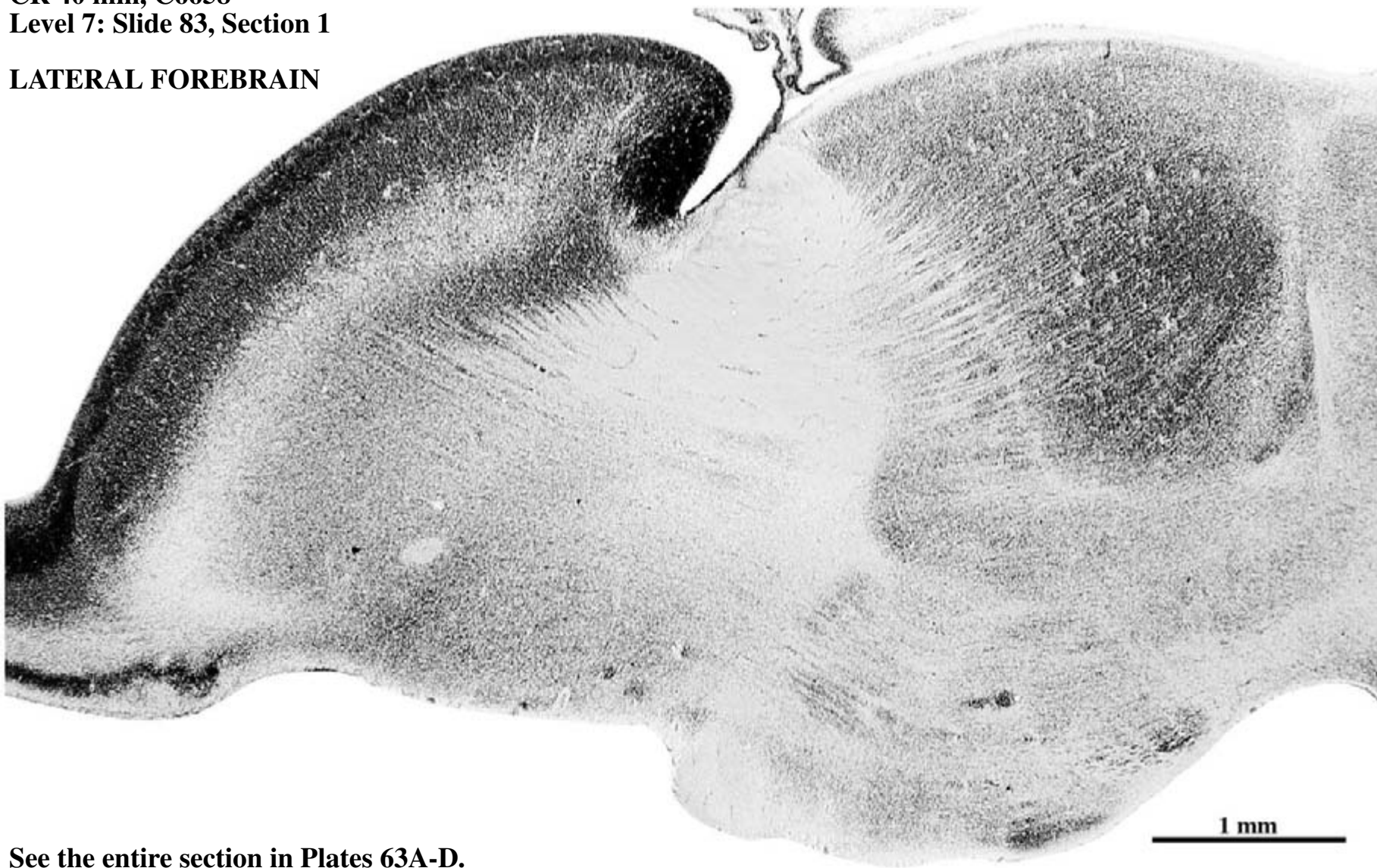


PLATE 73A

**GW9 Sagittal
CR 40 mm, C6658
Level 7: Slide 83, Section 1**

LATERAL FOREBRAIN



See the entire section in Plates 63A-D.

PLATE 73B

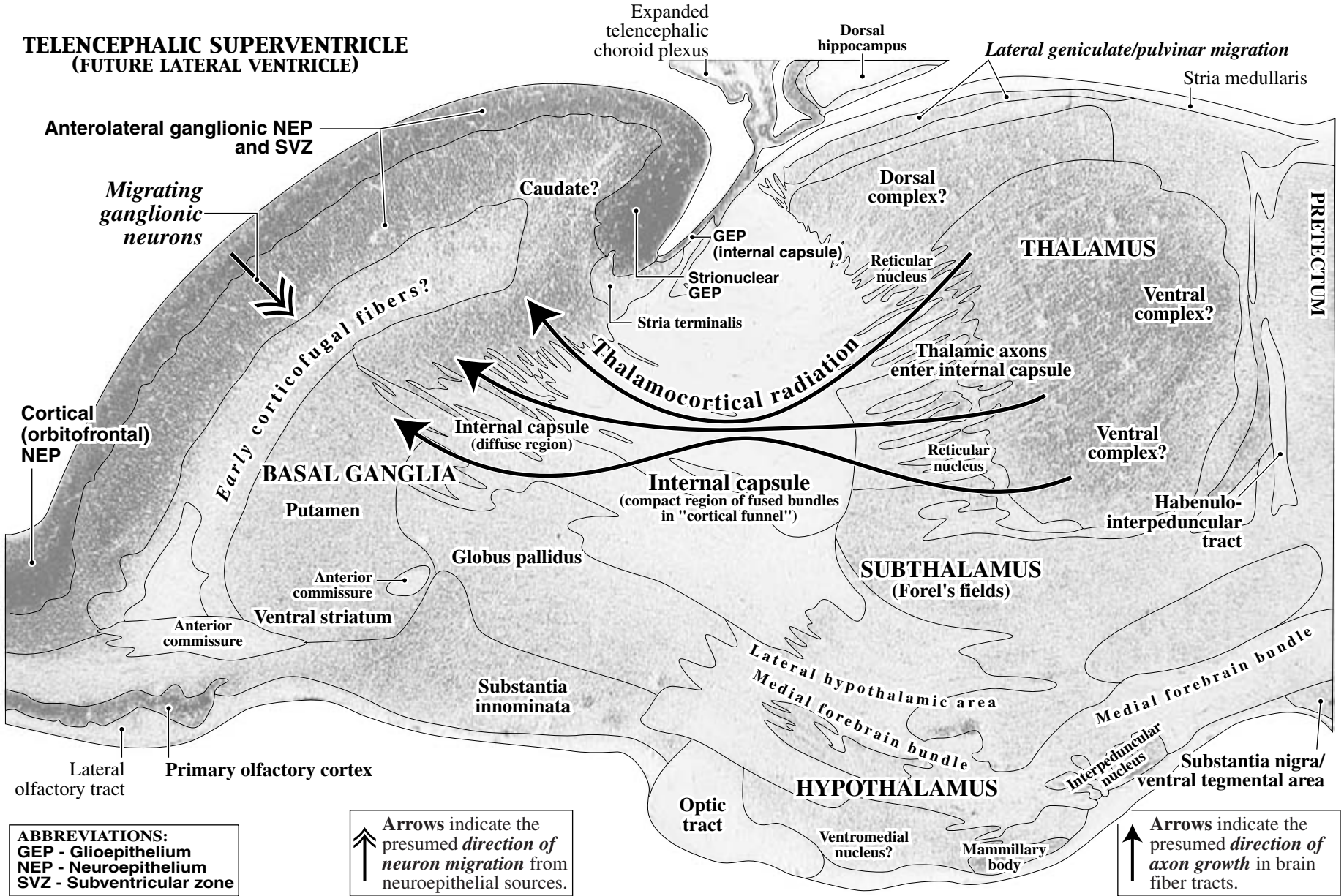
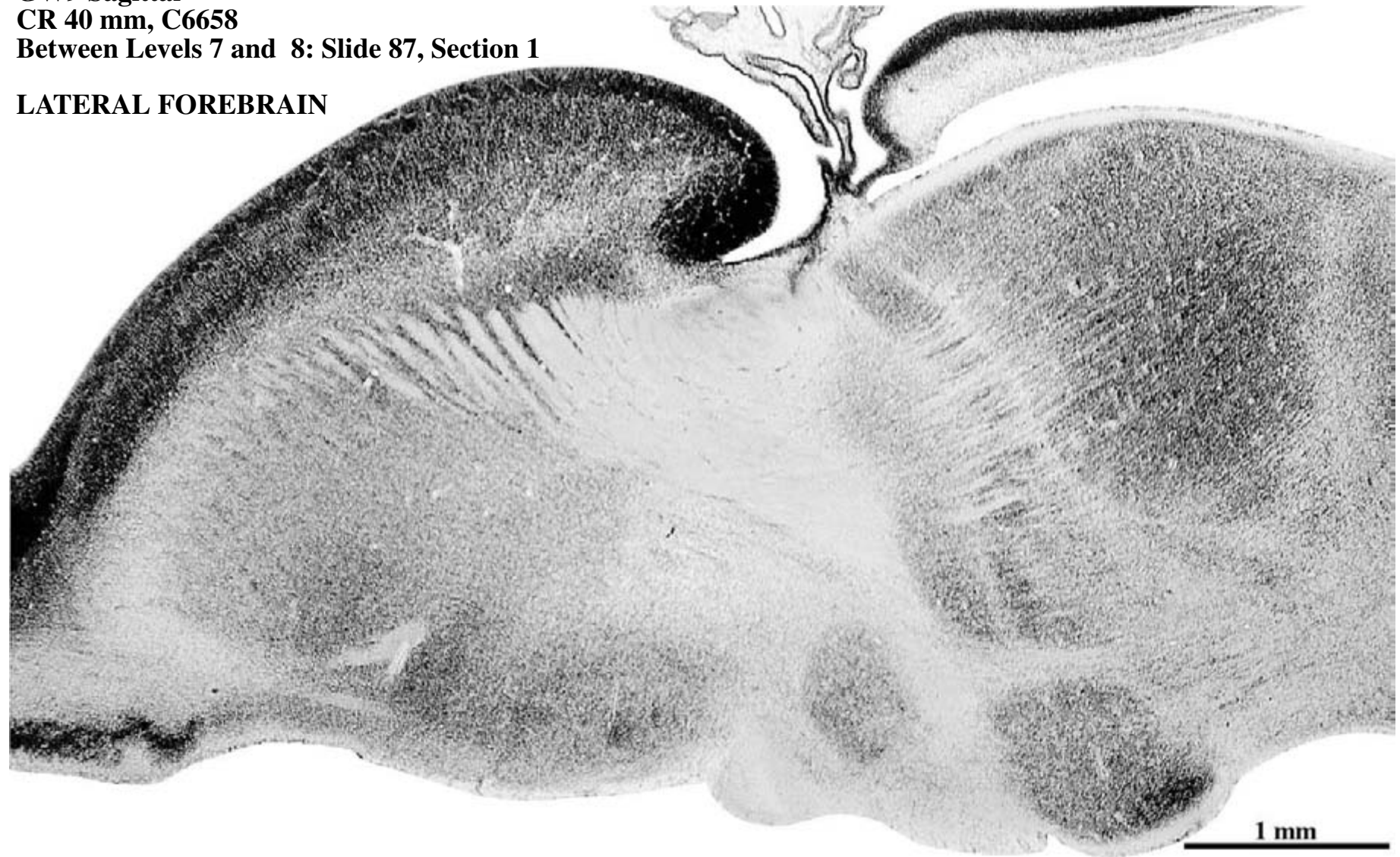


PLATE 74A

**GW9 Sagittal
CR 40 mm, C6658
Between Levels 7 and 8: Slide 87, Section 1**

LATERAL FOREBRAIN



See a low-magnification view of Level 7 in Plates 63A-D, Level 8 in Plates 64A-D.

PLATE 74B

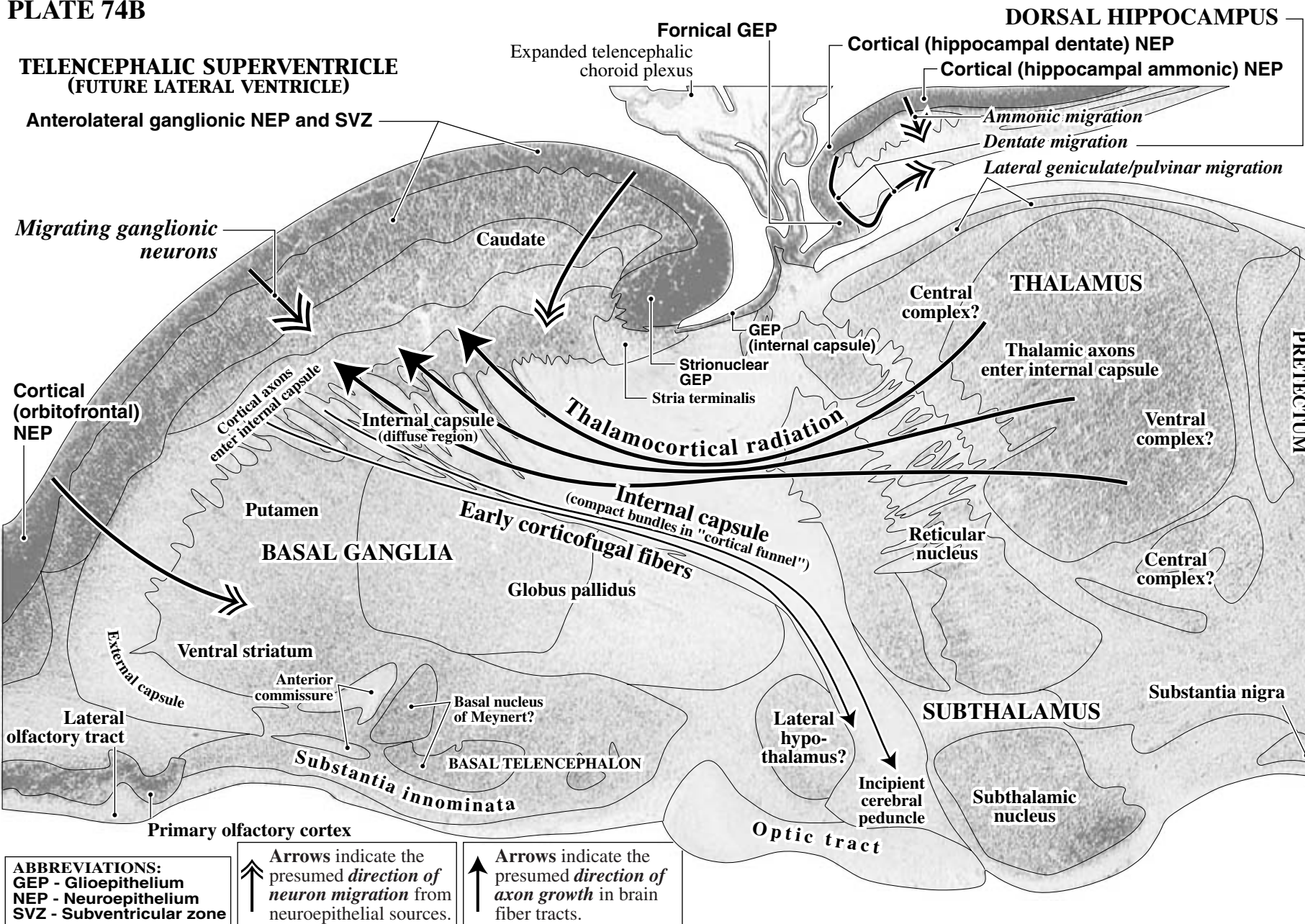
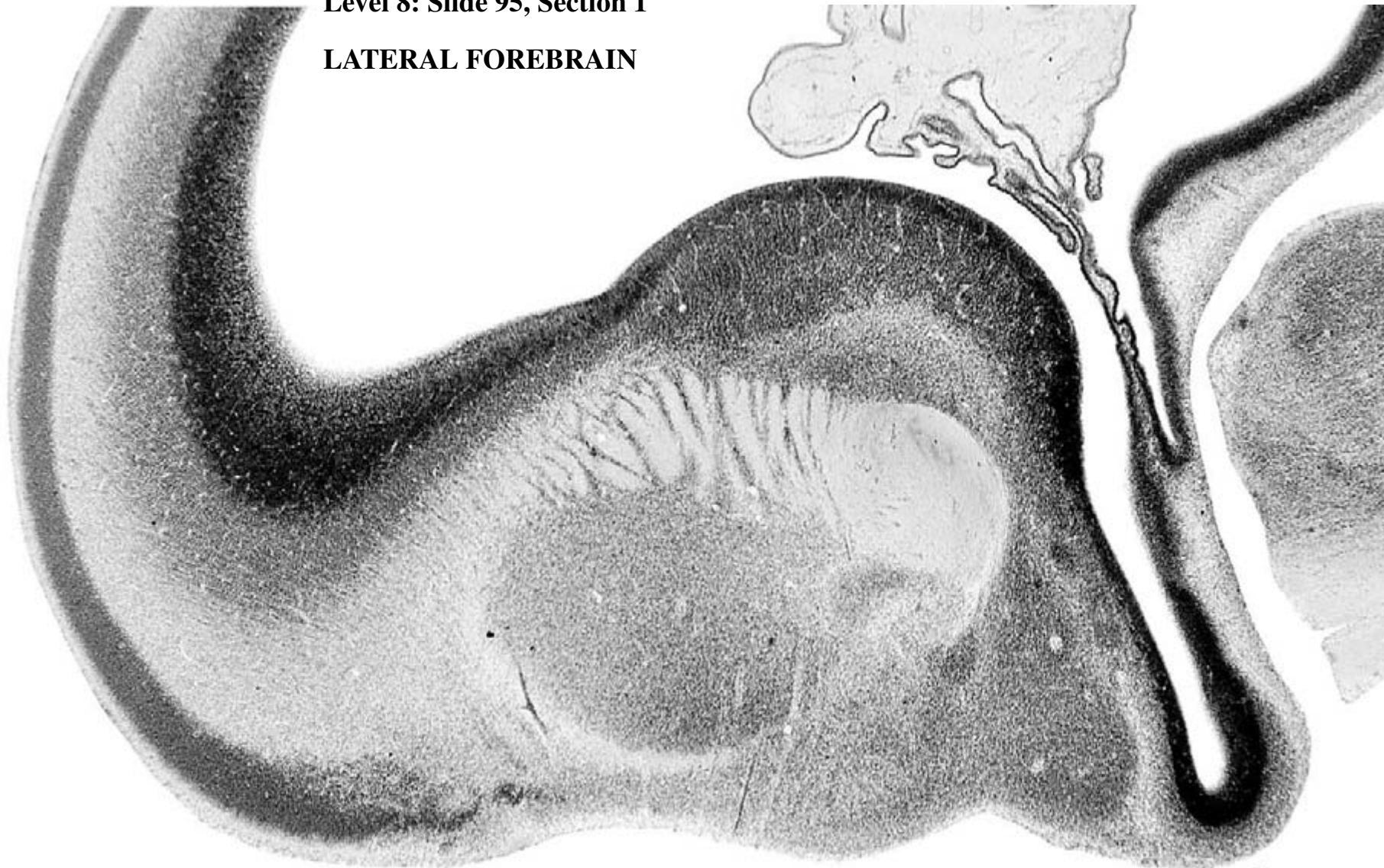


PLATE 75A

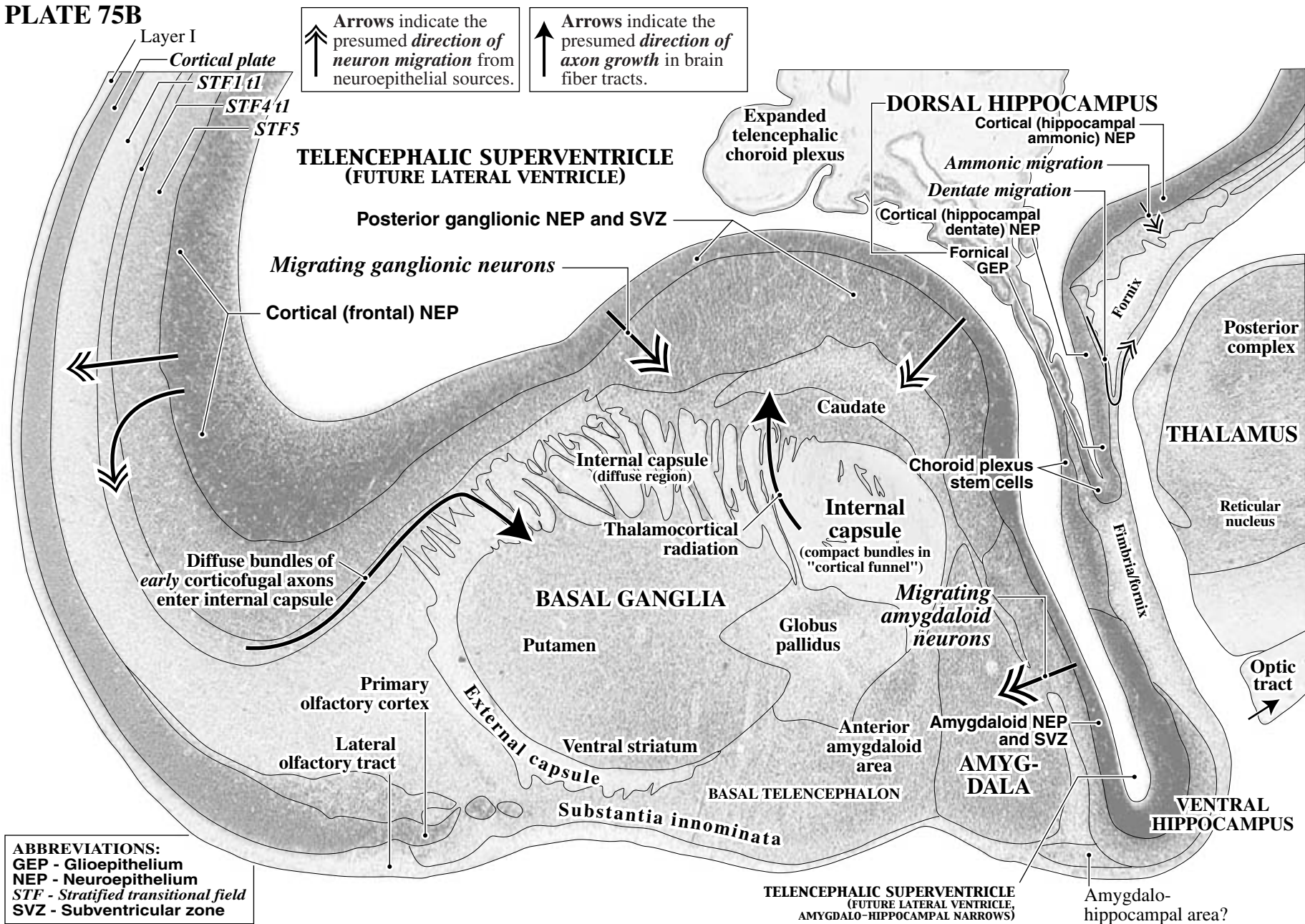
**GW9 Sagittal
CR 40 mm, C6658
Level 8: Slide 95, Section 1**

LATERAL FOREBRAIN



See a low-magnification view of Level 8 in Plates 64A-D.

PLATE 75B

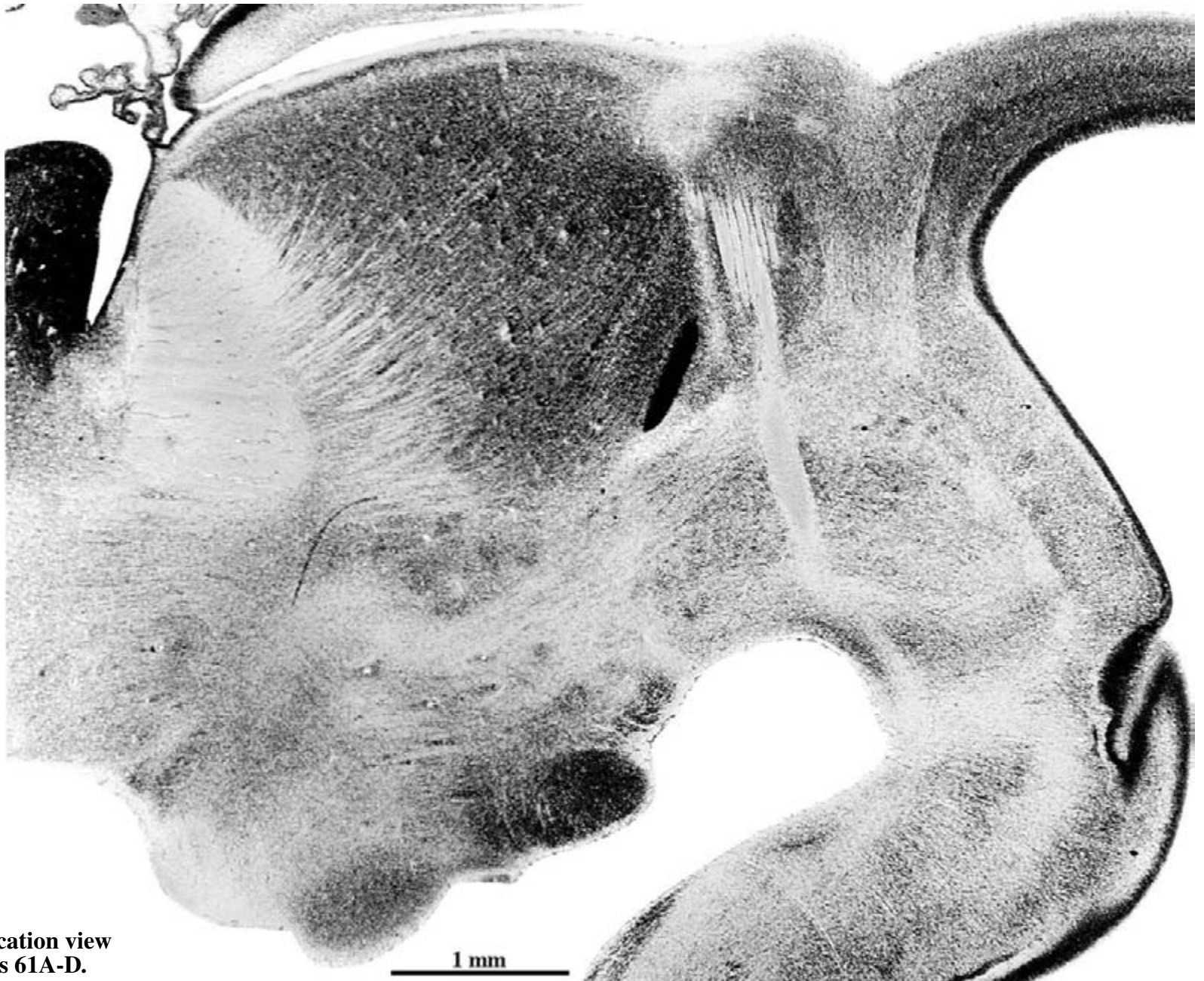


ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field
 SVZ - Subventricular zone

PLATE 76A

**GW9 Sagittal
CR 40 mm
C6658
Level 5:
Slide 79,
Section 2**

**DIEN-
CEPHALON
AND
MIDBRAIN**



**See a low-magnification view
of Level 5 in Plates 61A-D.**

PLATE 76B

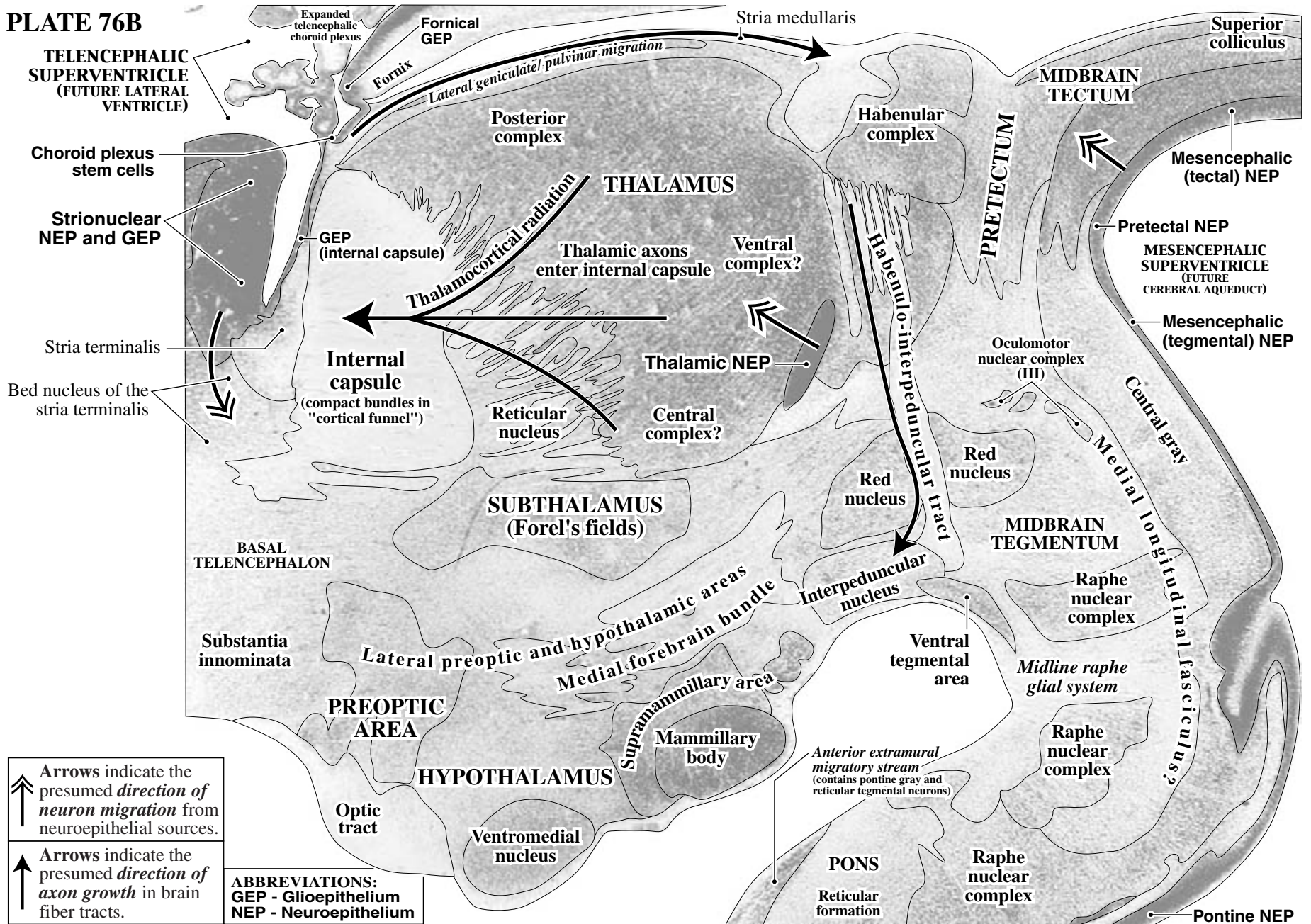


PLATE 77A

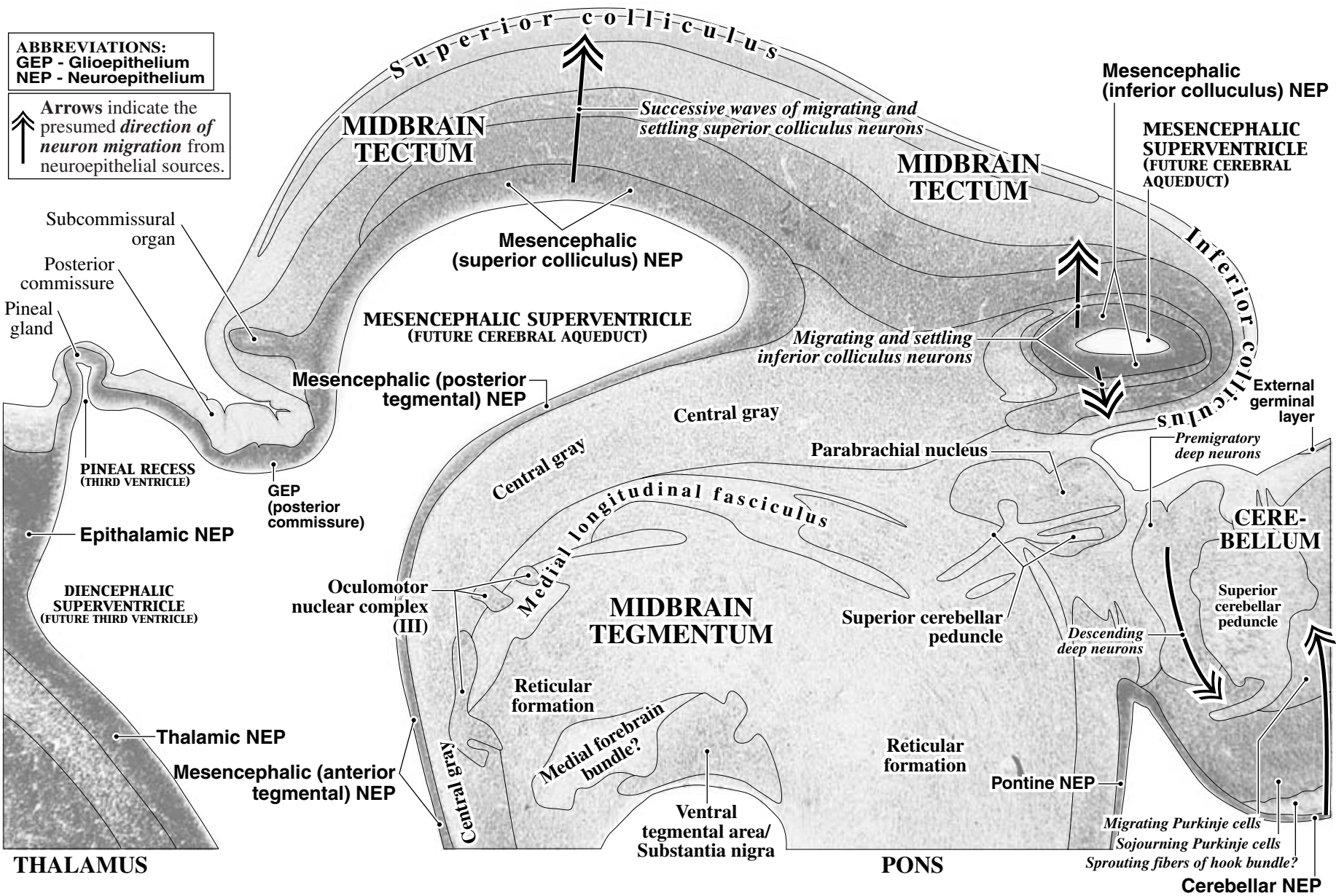
**GW9 Sagittal
CR 40 mm, C6658
Level 3:
Slide 67, Section 1**

MIDBRAIN



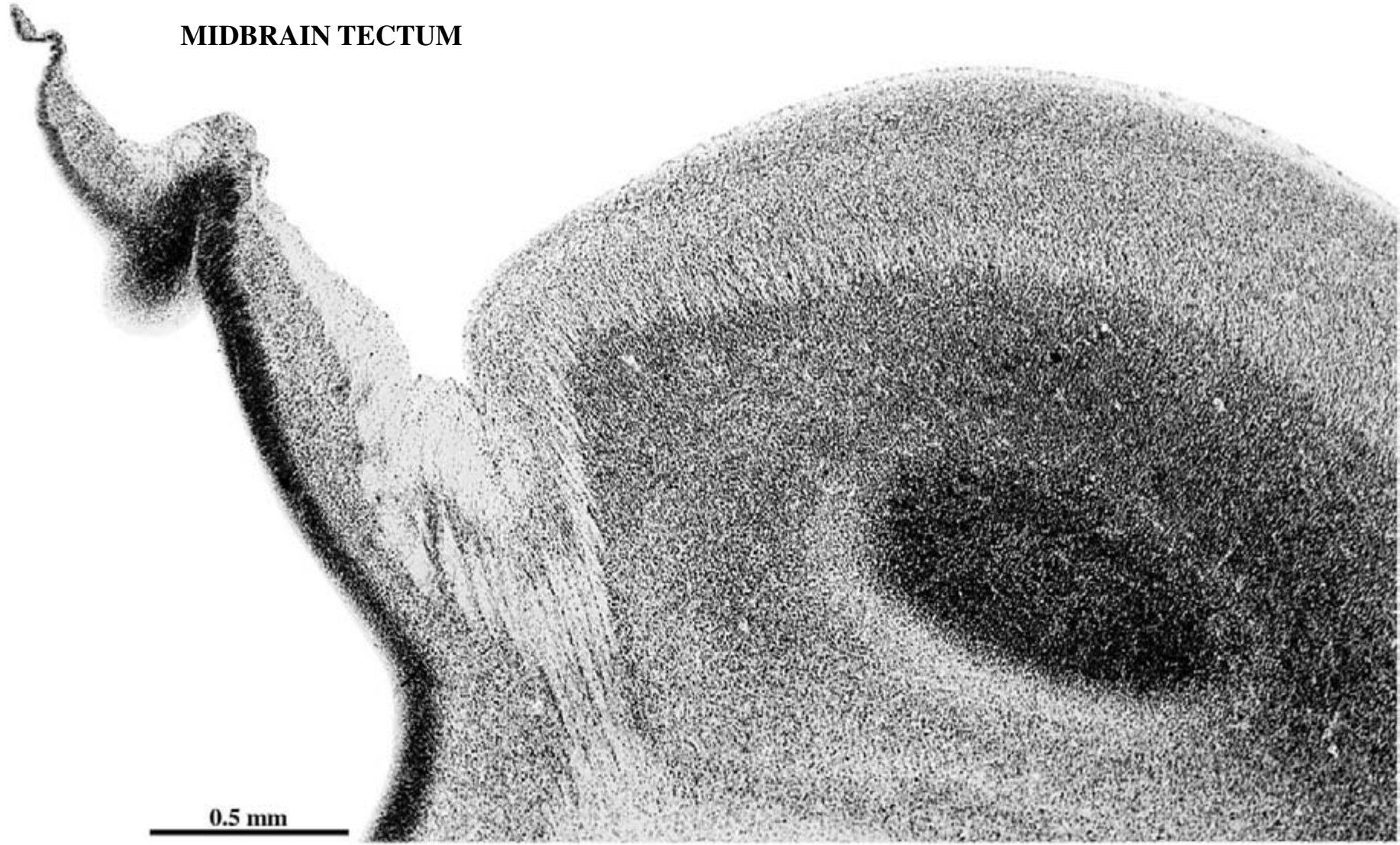
See a low-magnification view of Level 3 in Plates 59A-D.

PLATE 77B



**PLATE 78A GW9 Sagittal
CR 40 mm, C6658
Level 2: Slide 63, Section 1**

MIDBRAIN TECTUM



See a low-magnification view of Level 2 in Plates 58A-D.

PLATE 78B

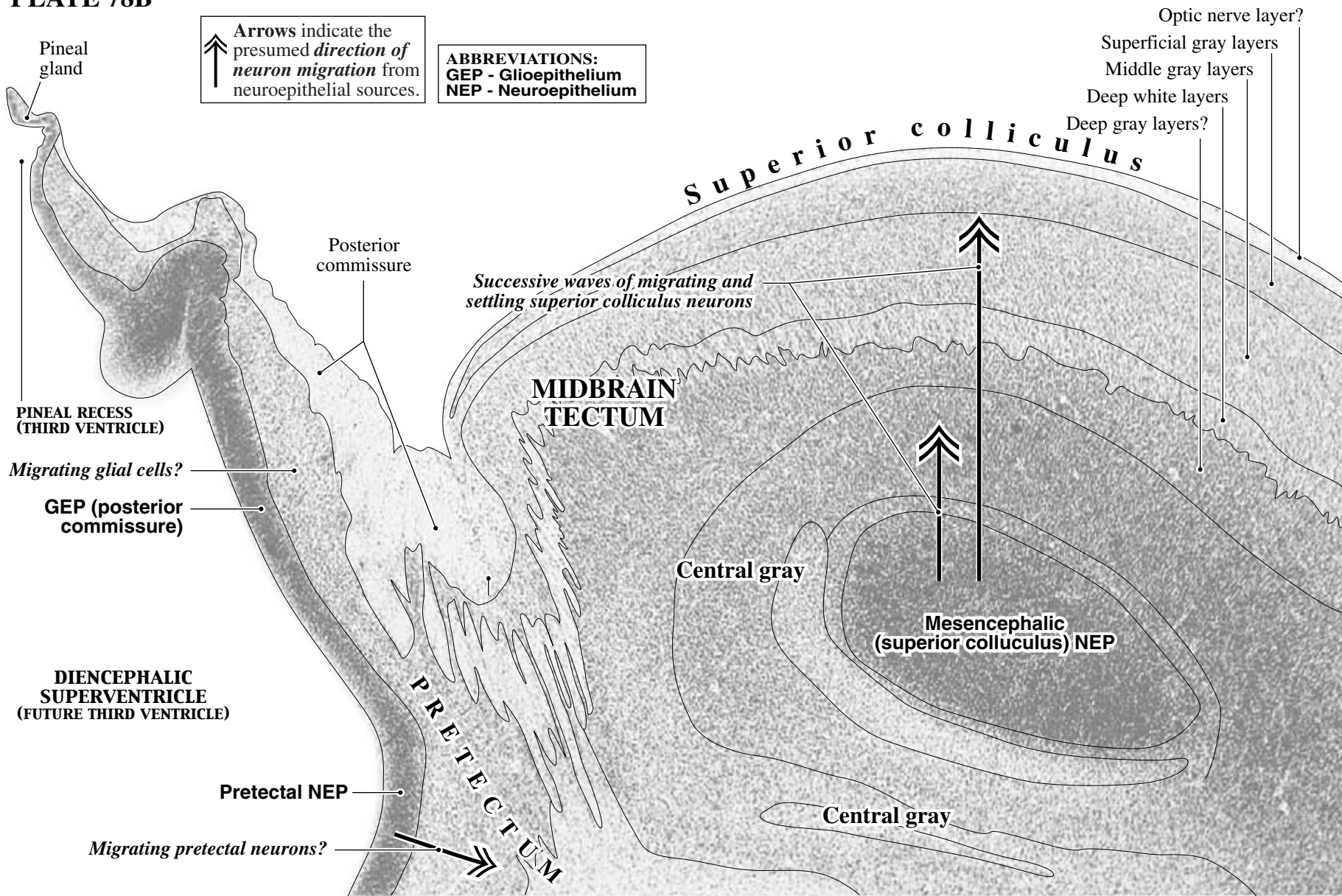
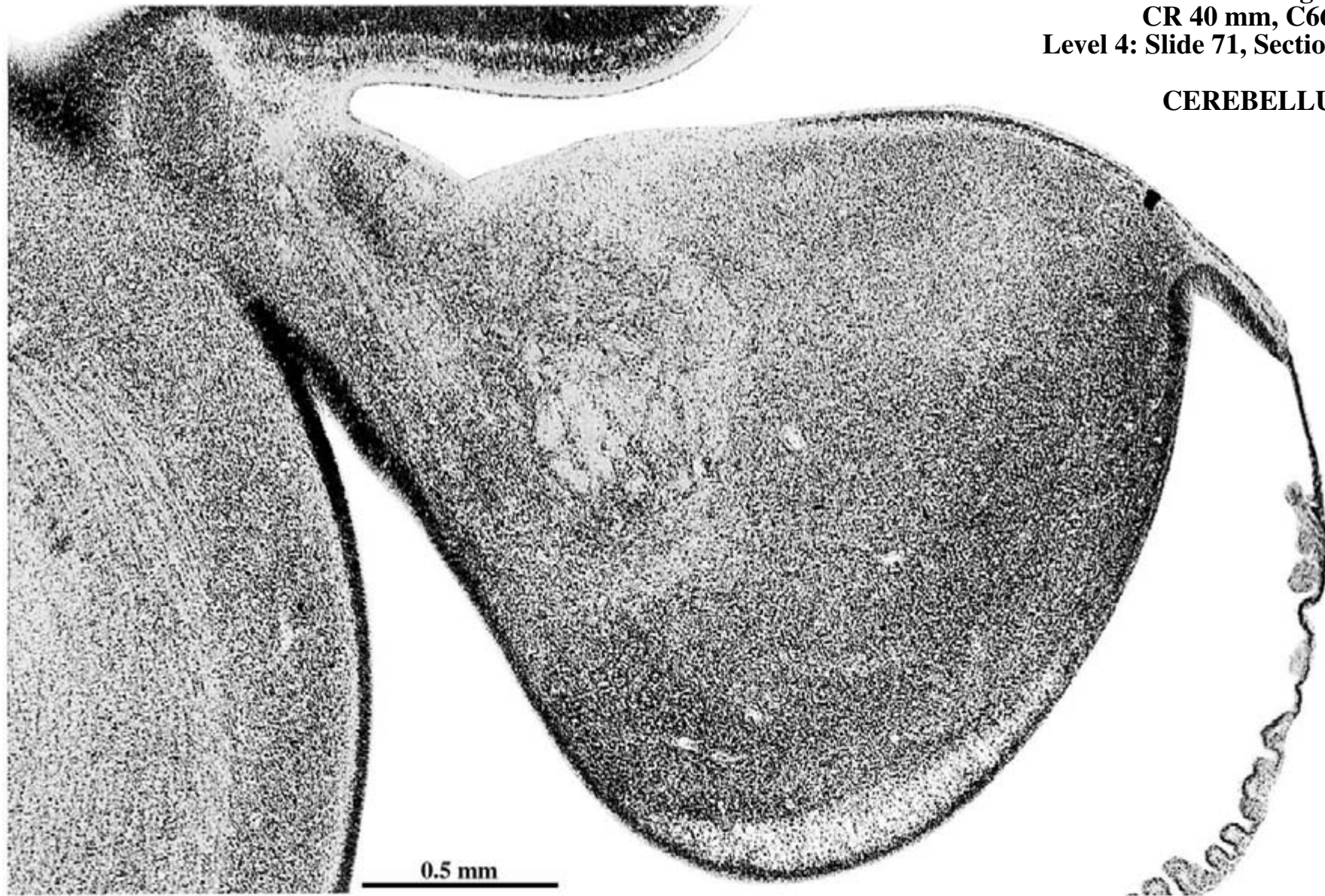


PLATE 79A

**GW9 Sagittal
CR 40 mm, C6658
Level 4: Slide 71, Section 2**

CEREBELLUM



See a low-magnification view of Level 4 in Plates 60A-D.

PLATE 79B

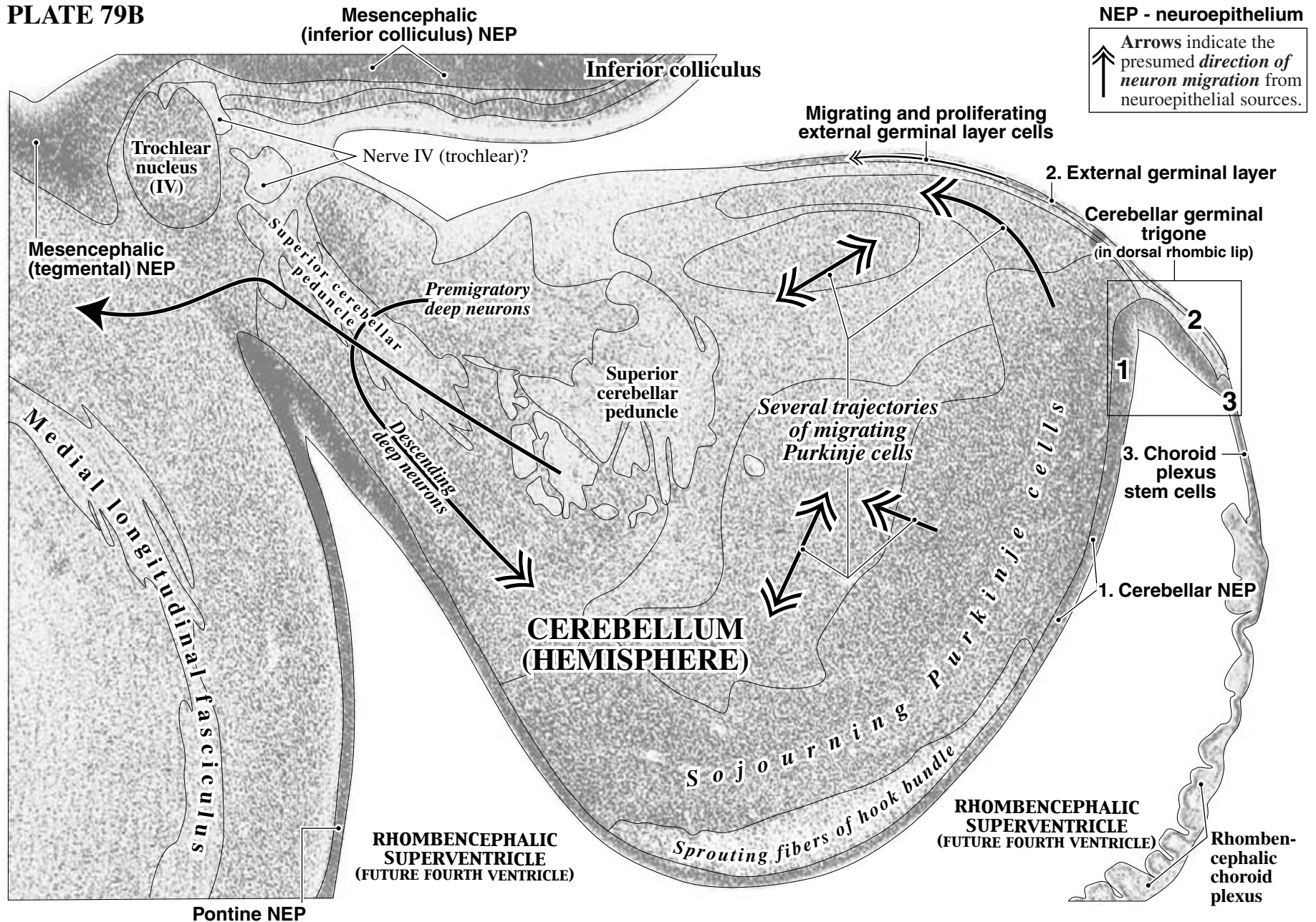
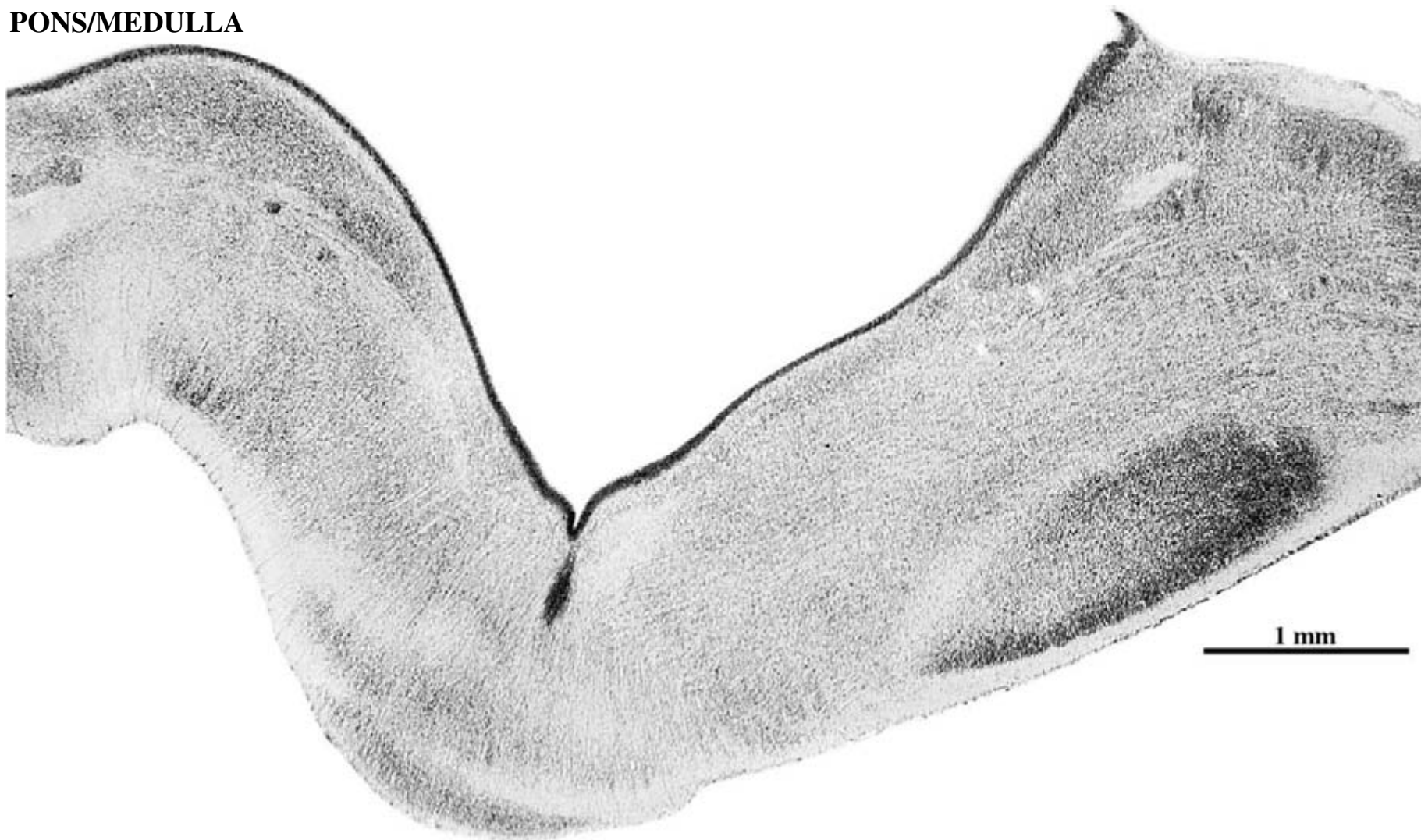


PLATE 80A

**GW9 Sagittal, CR 40 mm, C6658
Level 5: Slide 75, Section 2**

PONS/MEDULLA

See a low-magnification view of Level 5 in Plates 61A-D.

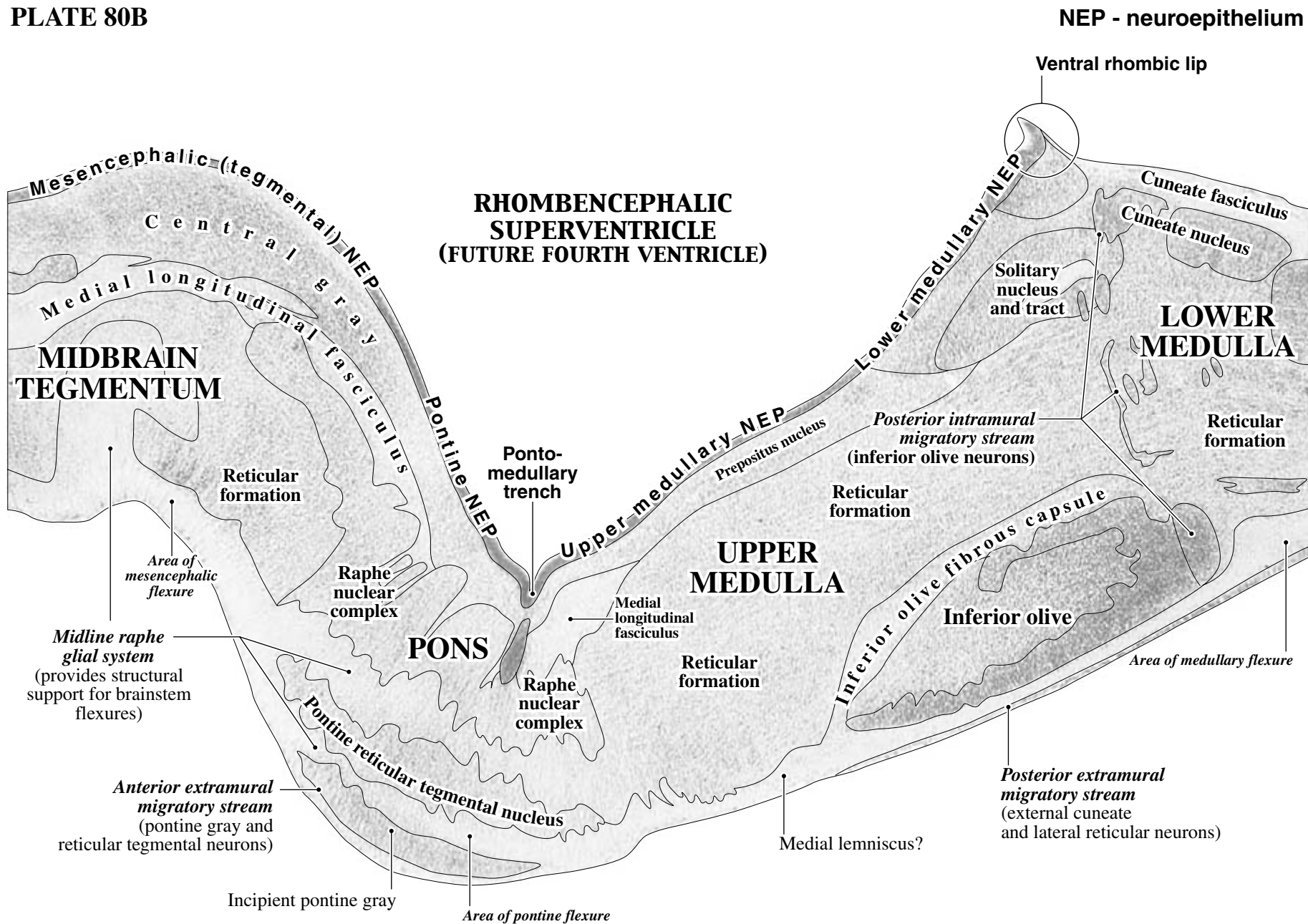
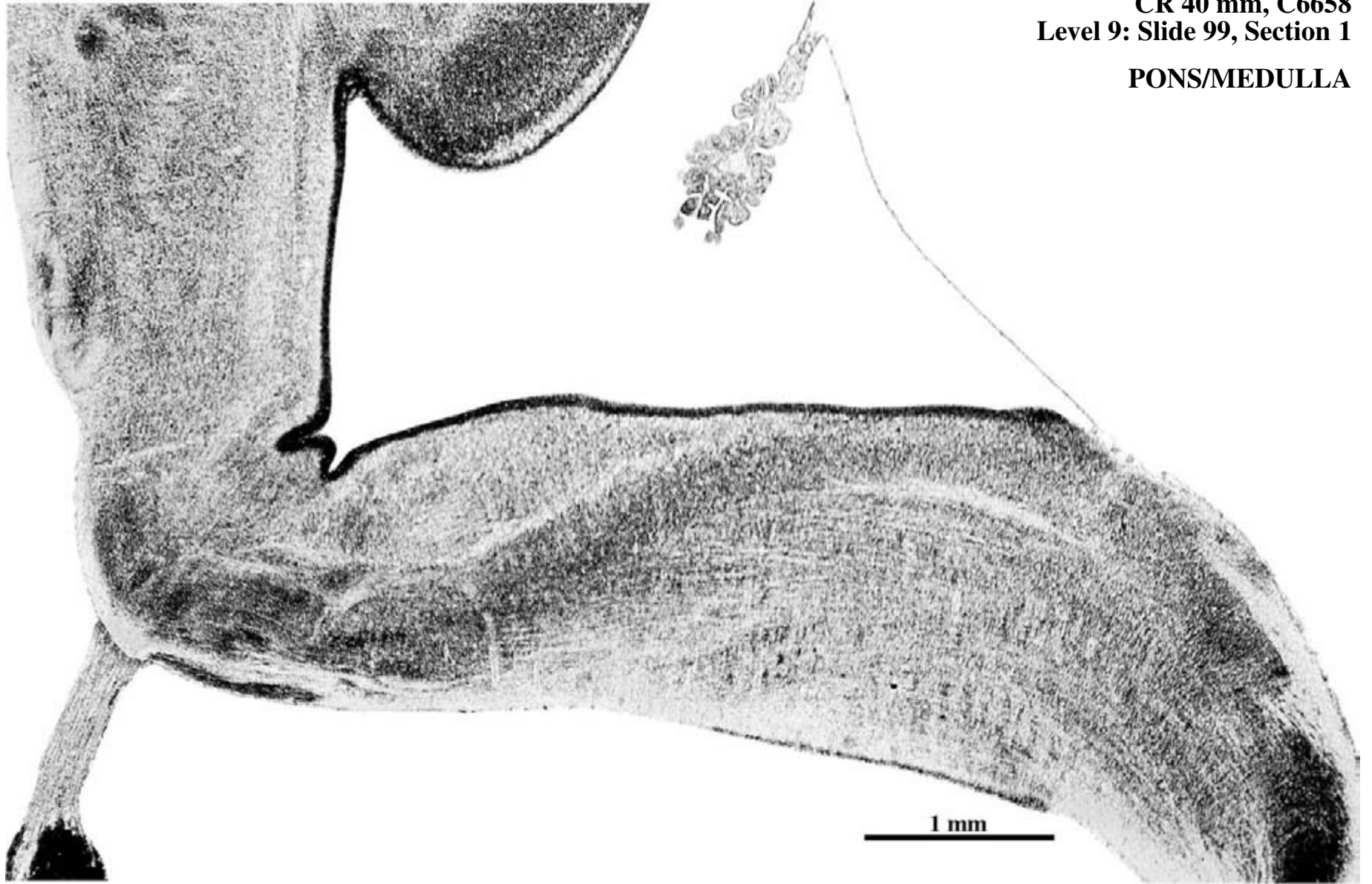


PLATE 81A

**GW9 Sagittal
CR 40 mm, C6658
Level 9: Slide 99, Section 1
PONS/MEDULLA**



See a low-magnification view of Level 9 in Plates 65A-D.

PLATE 81B

NEP - neuroepithelium

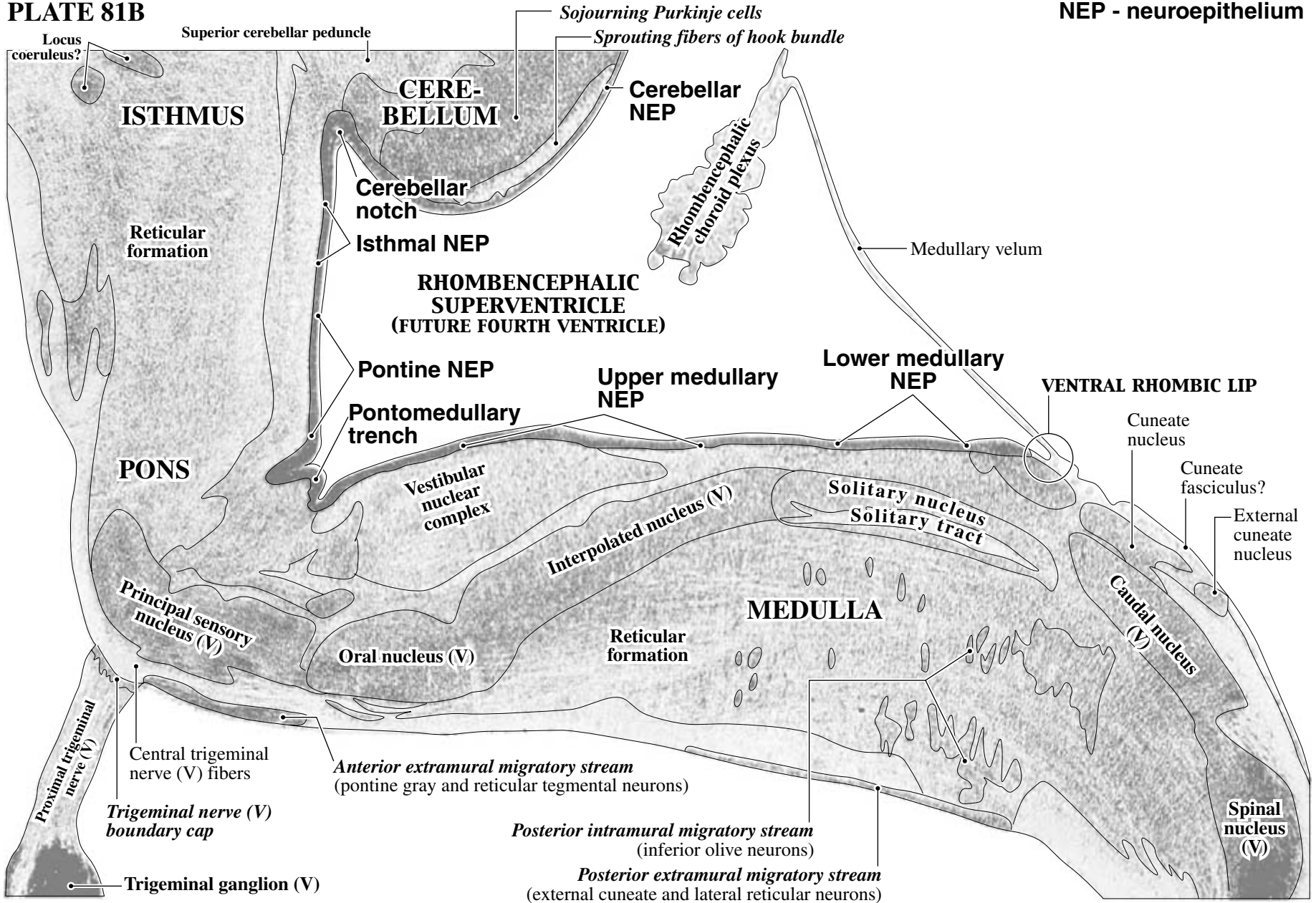
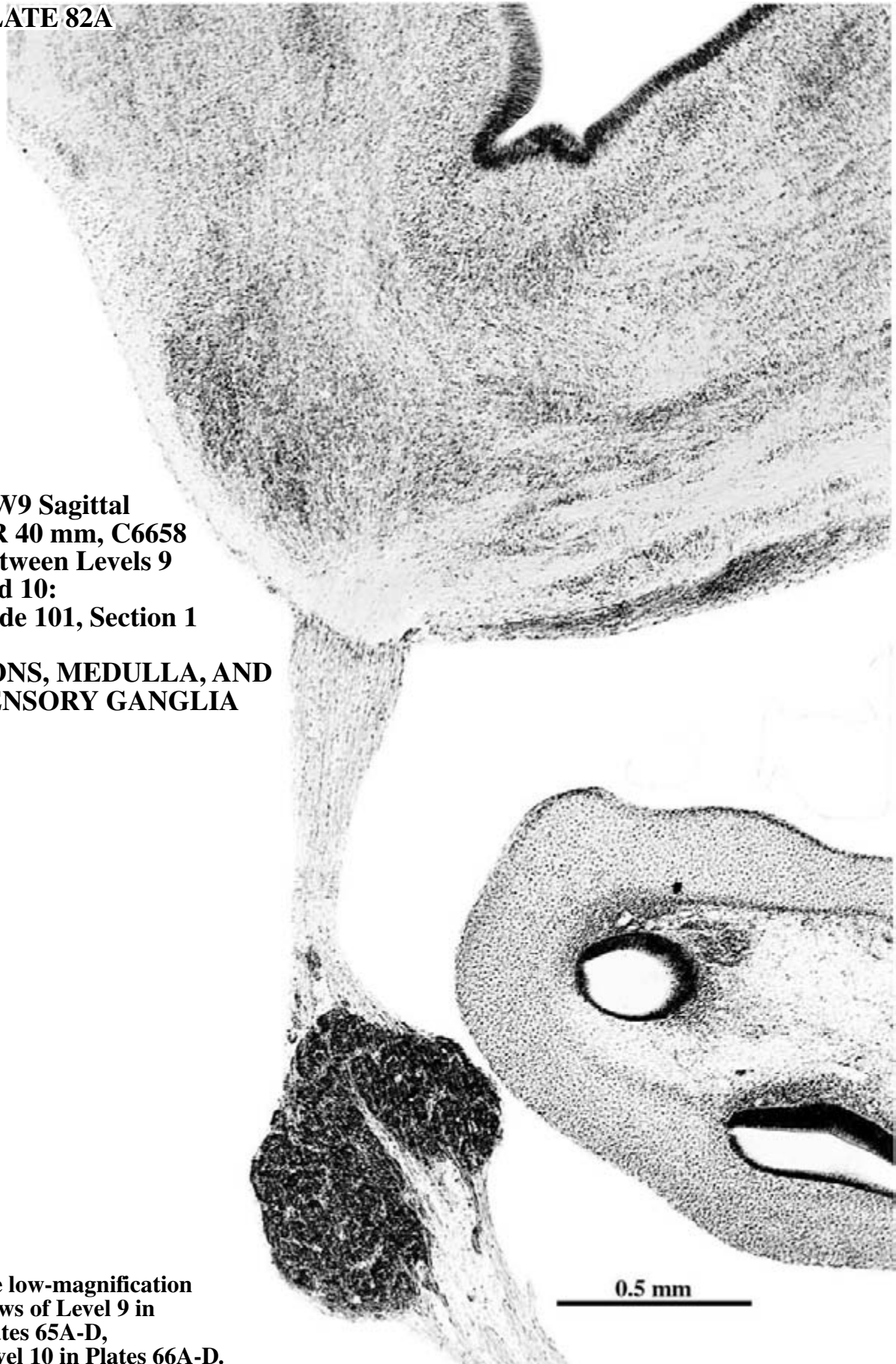


PLATE 82A

**GW9 Sagittal
CR 40 mm, C6658
Between Levels 9
and 10:
Slide 101, Section 1**

**PONS, MEDULLA, AND
SENSORY GANGLIA**

**See low-magnification
views of Level 9 in
Plates 65A-D,
Level 10 in Plates 66A-D.**



0.5 mm

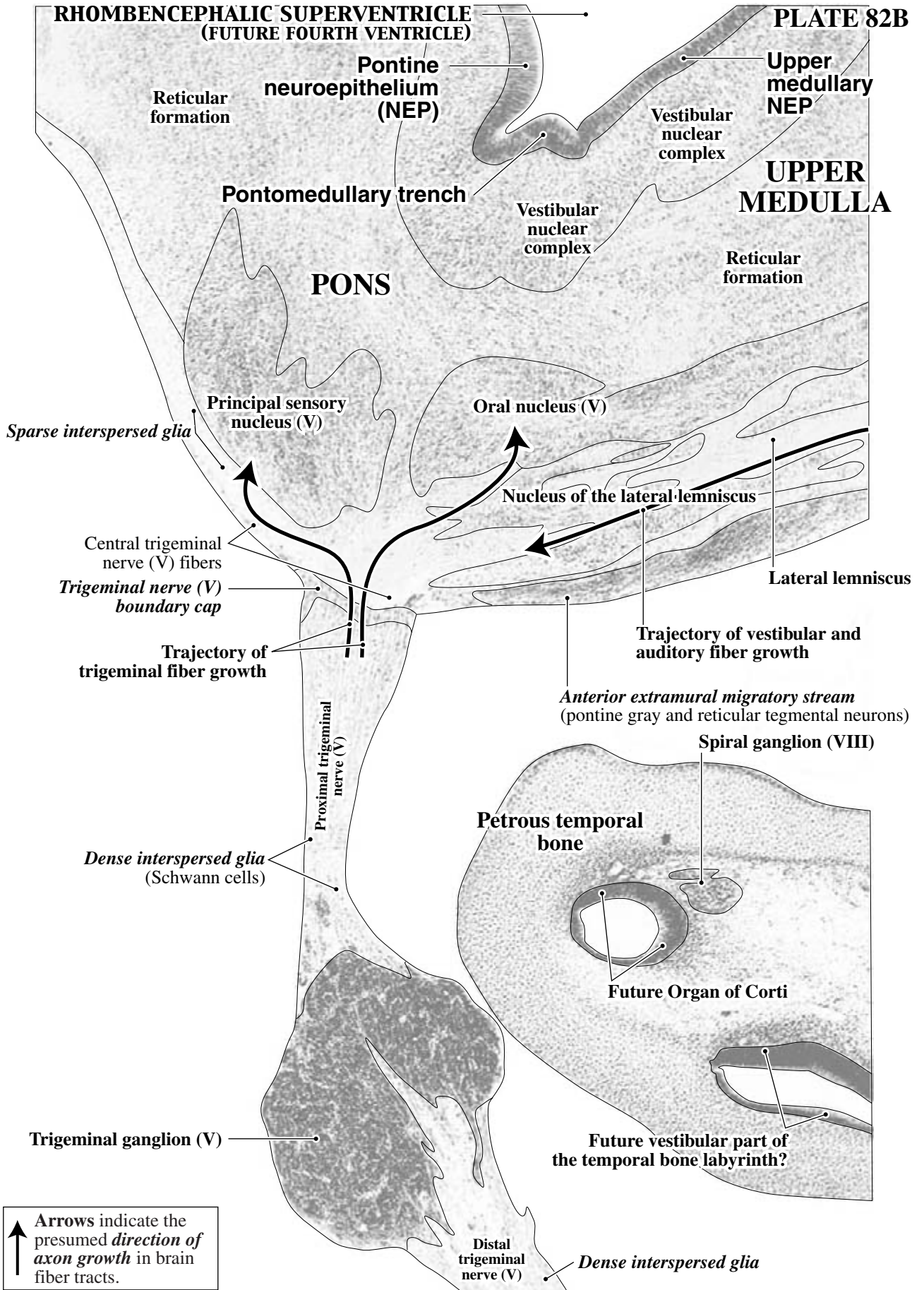
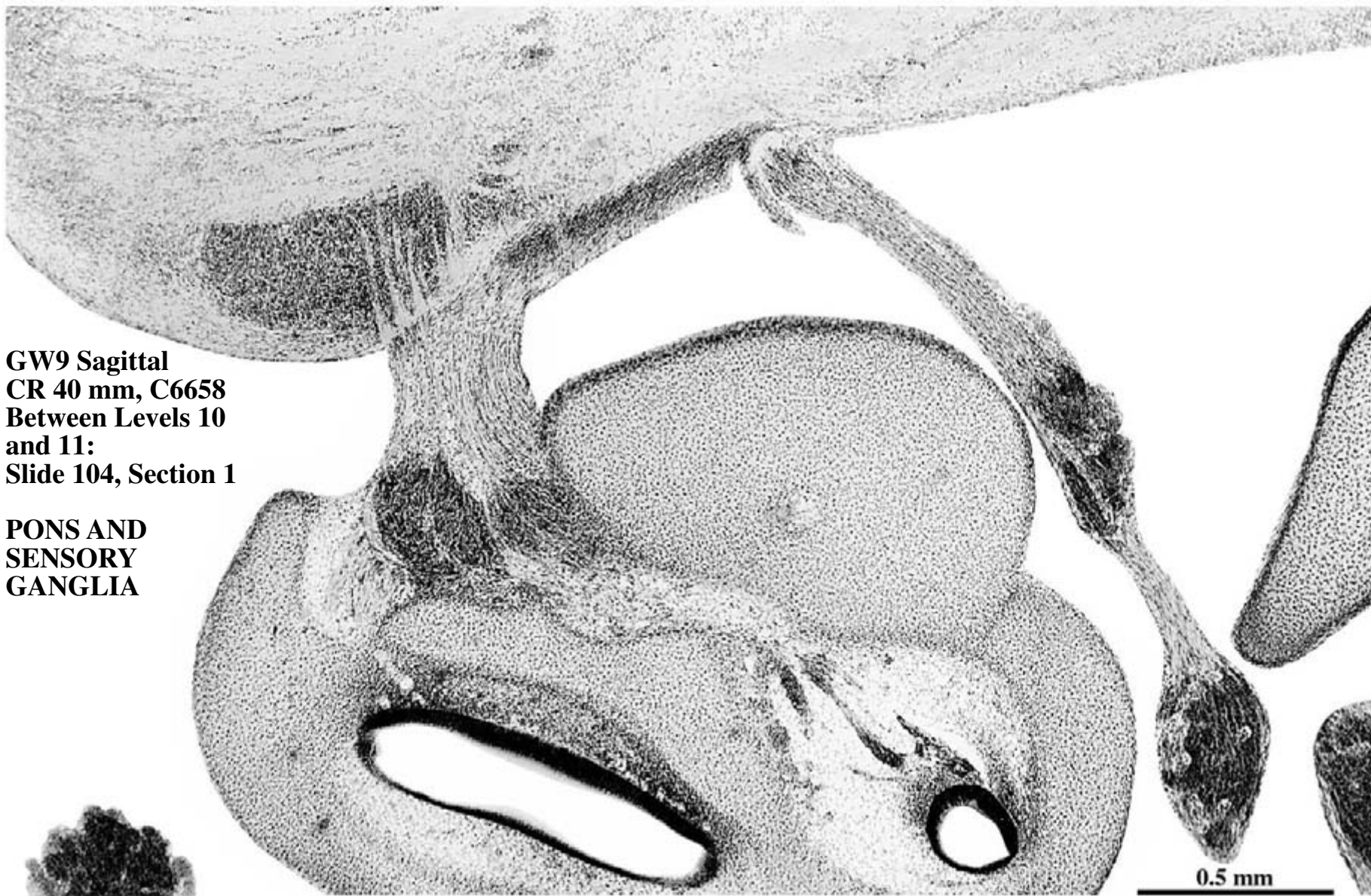


PLATE 83A

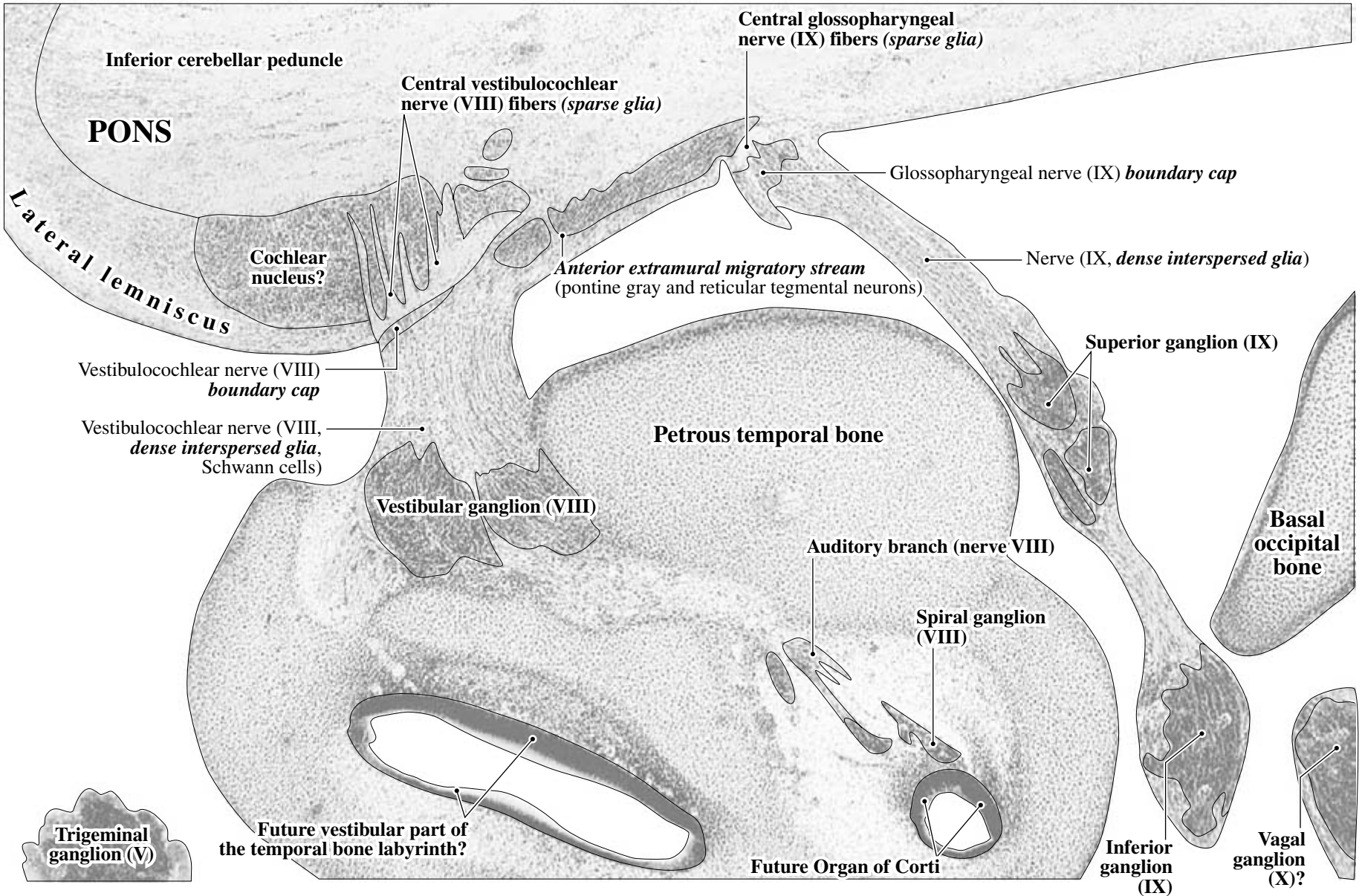
**GW9 Sagittal
CR 40 mm, C6658
Between Levels 10
and 11:
Slide 104, Section 1**

**PONS AND
SENSORY
GANGLIA**



See low-magnification views of Level 10 in Plates 66A-D, Level 11 in Plates 67A-D.

PLATE 83B



PART V: GW9 HORIZONTAL

This is specimen number 886 in the Carnegie Collection, designated here as C886. A normal fetus with a crown-rump length (CR) of 43 mm was collected in 1914 after a hysterectomy due to pelvic inflammation. The fetus is estimated to be at gestational week (GW) 9. The entire fetus was fixed in Bouin's, embedded in celloidin, and 100 μm sections were cut in a plane midway between the horizontal and coronal planes (**Figure 4**). Because many of the sections do not contain the cerebral cortex and the brainstem is cut in a crosswise direction, it more closely resembles a horizontally sectioned brain. All sections were stained with hematoxylin and eosin. Since there is no photograph of C886's brain before it was embedded and cut, a specimen from Hochstetter (1919) is used to show the external features of the brain at GW9 (**Figure 4**). **Levels 1-11**, generally larger sections containing the cerebral hemispheres, are shown at low magnification in **Plates 84A/B-93A/B**. The core parts of the sections in **Levels 3-11** are also shown at high magnification in **Plates 85C/D-93C/D**. **Levels 12-19**, generally smaller sections containing only the brainstem, are shown at a high magnification in **Plates 94A/B-101A/B**. To more efficiently use page space, all plates are in landscape orientation (anterior/ventral: left side of photograph, bottom of page; posterior/dorsal: right side of photograph, top of page).

C886 is similar to C6658 in the level of brain maturation. The chief reason for including this specimen is to provide a different perspective for viewing brain structure at GW9. In the cerebral cortex, the *neuroepithelium* is prominent as the sole germinal matrix; the *stratified transitional field (STF)* consists of *STF1*, *STF5*, and *STF4* only in lateral areas. The anterolateral (thicker) to dorso-medial (thinner) maturation gradient in the cortical plate and STF layers are evident. In anterolateral parts of the cerebral cortex, streams of neurons and glia appear to leave *STF4* and enter the *lateral migratory stream*. The hippocampus contains *ammonic and dentate migrations*, but there is no evidence of a pyramidal in Ammon's horn or a dentate gyrus. A massive *neuroepithelium/subventricular zone* overlies the amygdala, nucleus accumbens, and stria-

tum (caudate and putamen) where neurons (and glia) are being generated.

The cerebellum is a thick, smooth plate overlying the posterior pons and medulla, and a definite *neuroepithelium* at the ventricular surface, indicating some Purkinje cells are still being generated. Many Purkinje cells are sojourning in a dense layer outside the neuroepithelium, and others are migrating upward. Many of the deep neurons are superficial in the cerebellum, but some are migrating downward to intermingle with upwardly migrating Purkinje cells. The cortical surface is partially covered by an *external germinal layer (egl)* that is actively producing neuronal stem cells, as it grows over the surface of the cerebellar cortex.

The third ventricle, aqueduct, and fourth ventricle are lined by thin *neuroepithelia*. The midbrain tegmentum, pons, and medulla have the thinnest neuroepithelia indicating that only the latest generated neurons are being produced at this time. The thick precerebellar neuroepithelium is an exception in the medulla. Thicker neuroepithelia are in the cerebellum (see above) and midbrain tectum, indicating many neurons are still being generated, although the majority of the neurons in these sites are already post-mitotic. The neuroepithelium is still more thick in the hypothalamus and thalamus, in accordance with the later maturation of the diencephalon compared to the rest of the brainstem.

Neurons throughout the diencephalon, midbrain tegmentum, pons, and medulla are migrating and settling. This specimen shows a very prominent migration of subthalamic nucleus neurons from the posterior hypothalamic neuroepithelium. Except for the subthalamic nucleus, nuclear divisions are very indistinct throughout the diencephalon. More definition is seen in the midbrain tegmentum, pons, and medulla. The *anterior extramural* and *posterior extramural migratory streams* are dense subpial accumulations in the medulla and pons.

GW9 "HORIZONTAL" SECTION PLANES

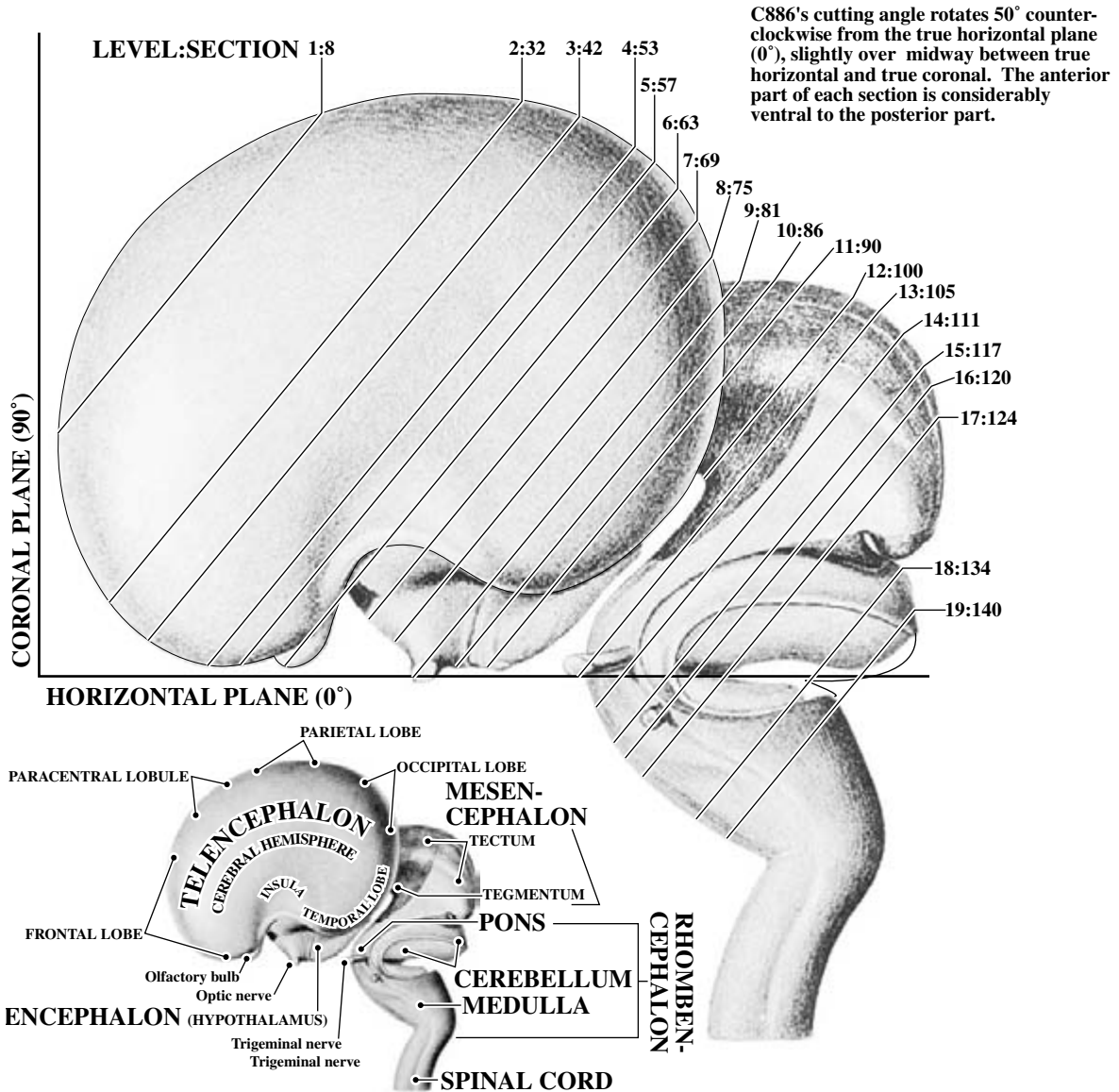


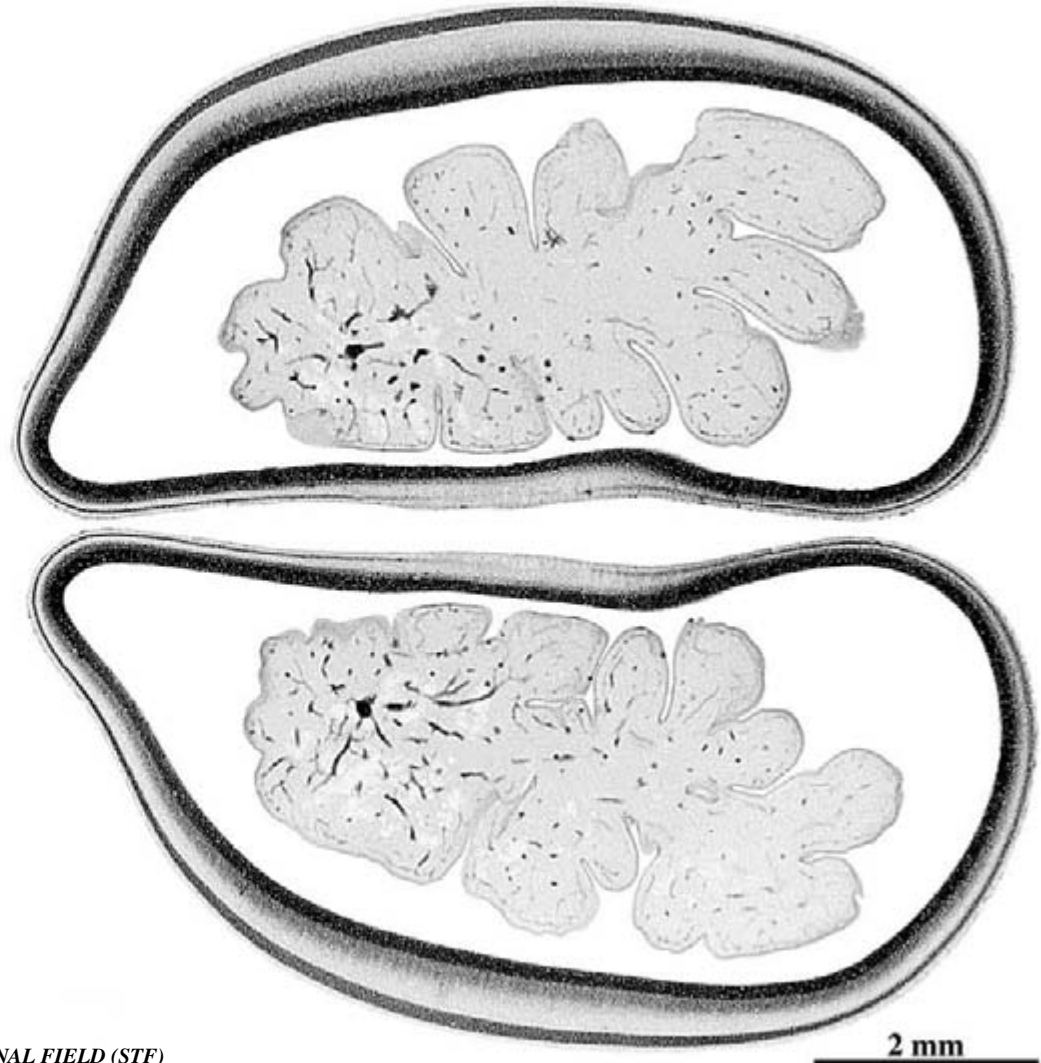
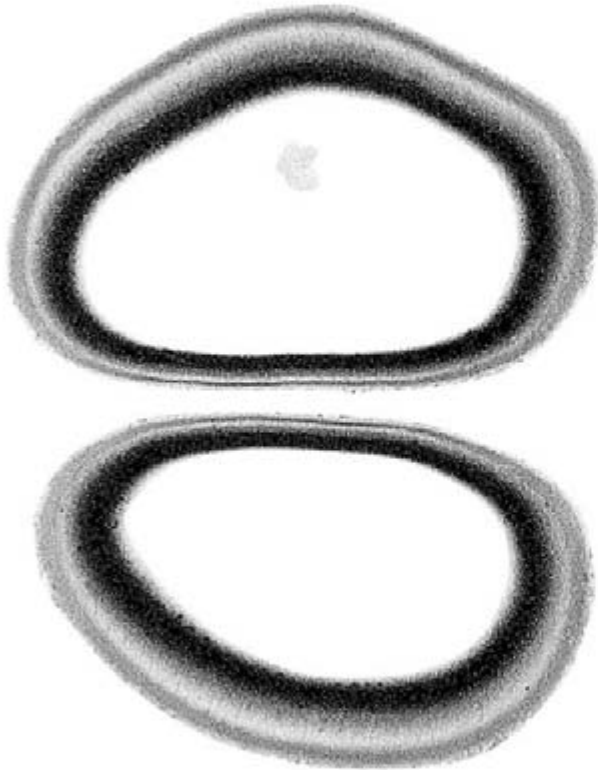
Figure 4. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 38 mm (modified from Figure 43, Table VII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of C886 in the following pages. The small inset identifies the major structural features. The line in the cerebellum and dorsal edges of the pons and medulla is the cut edge of the medullary velum.

PLATE 84A

GW9 Horizontal
CR 43 mm, C886

Level 2: Section 33

Level 1: Section 8

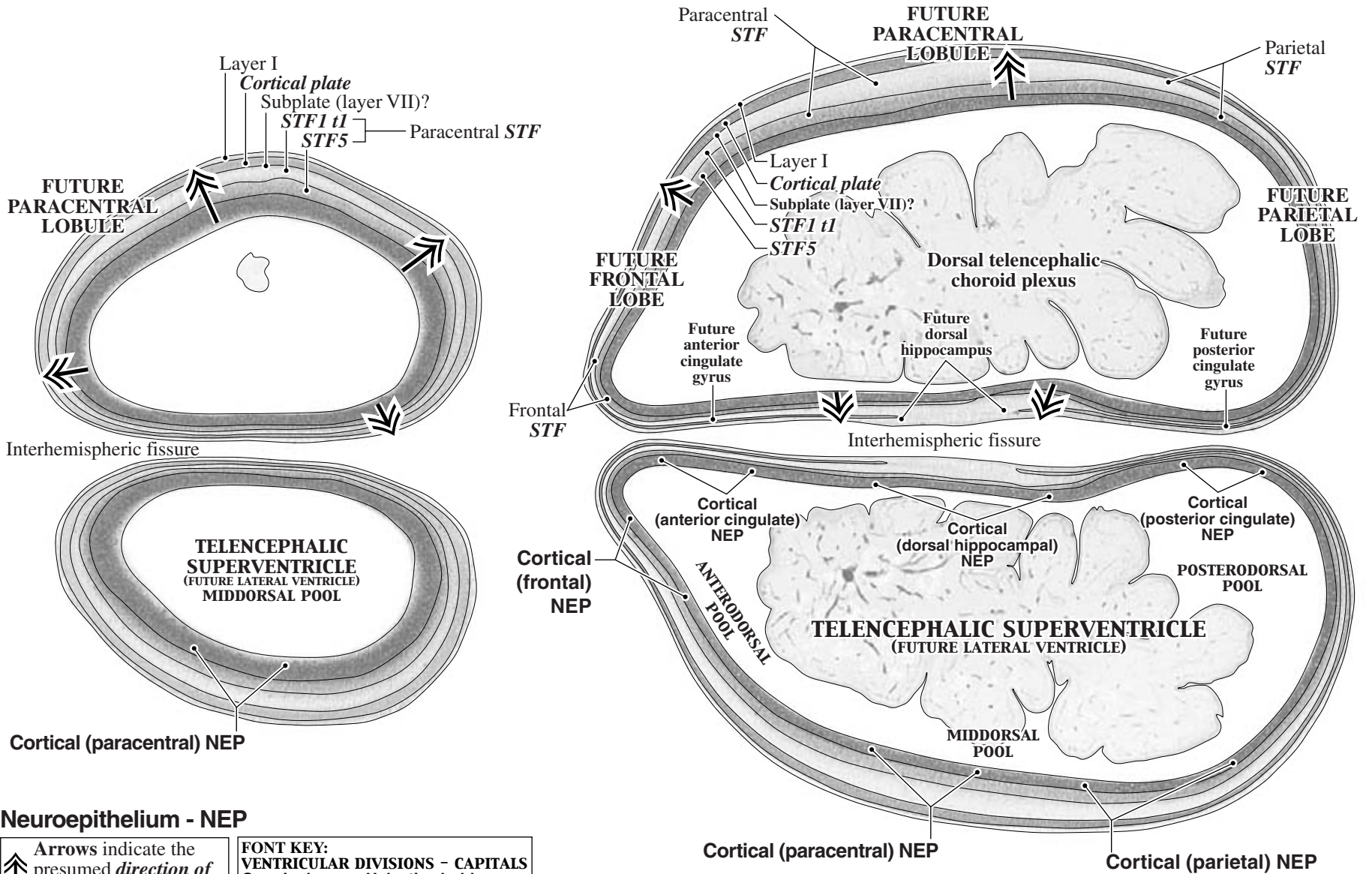


LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 84B



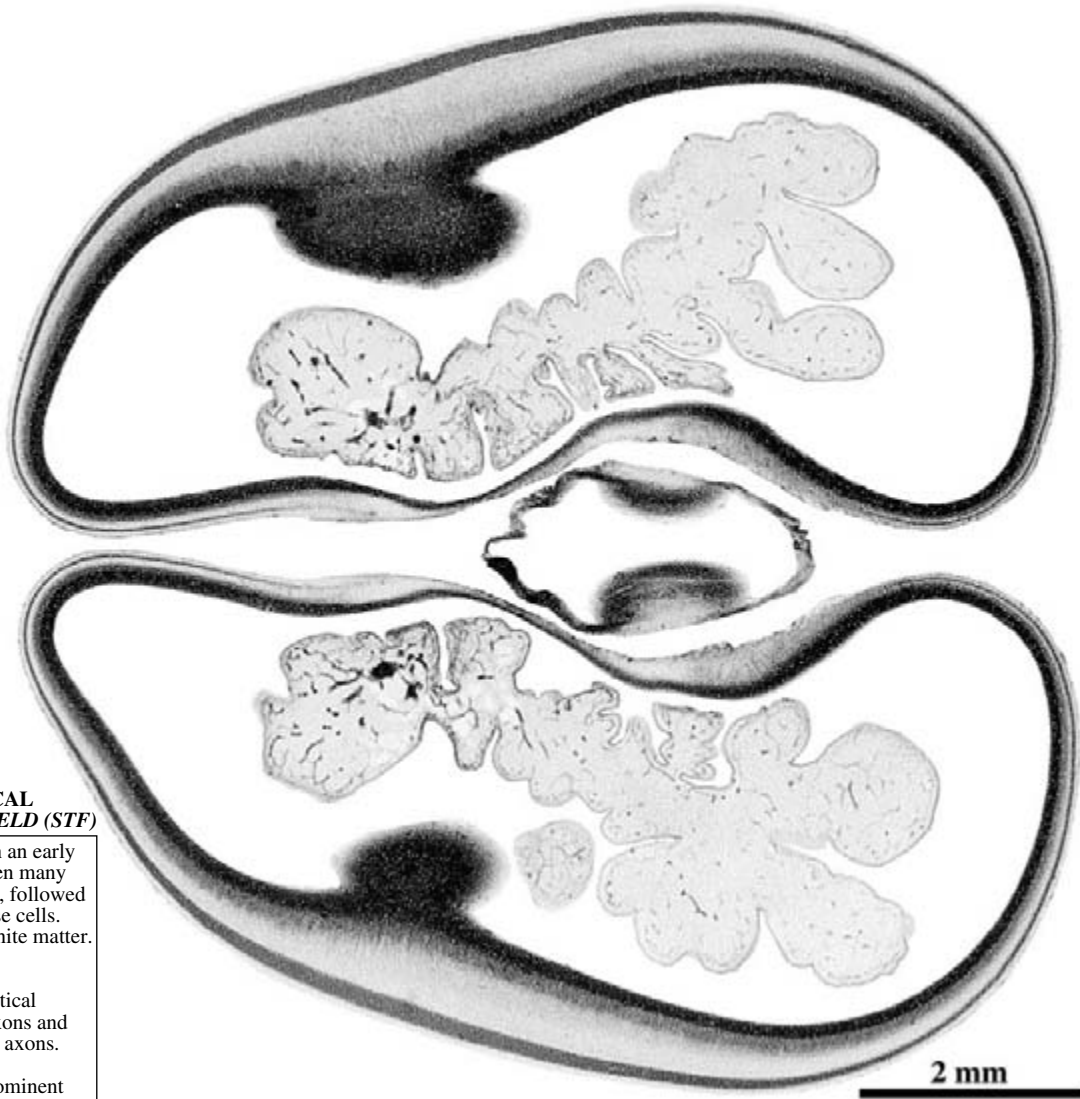
Neuroepithelium - NEP

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 85A

**GW9 Horizontal
CR 43 mm, C886
Level 3: Section 42**



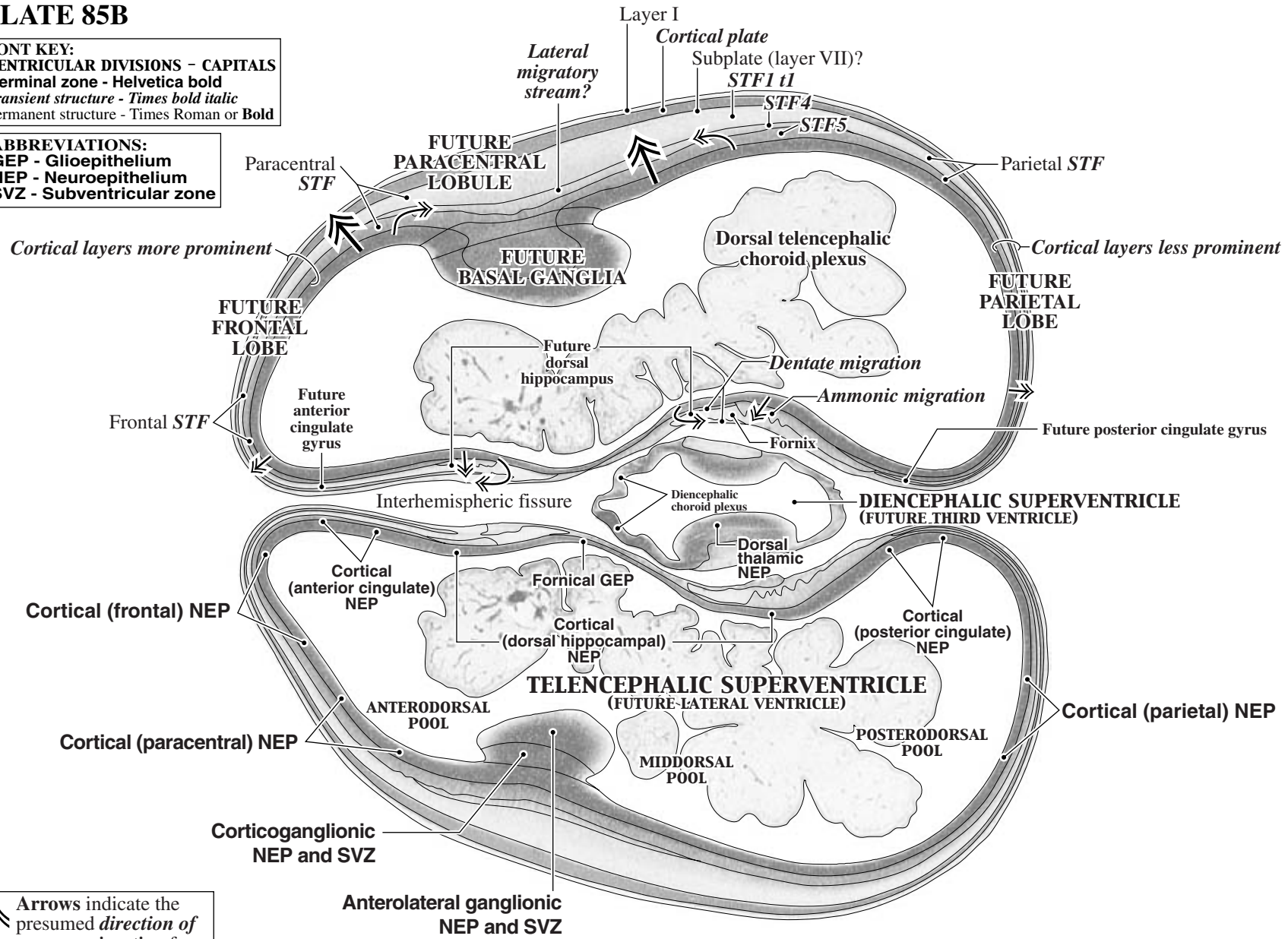
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 85B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

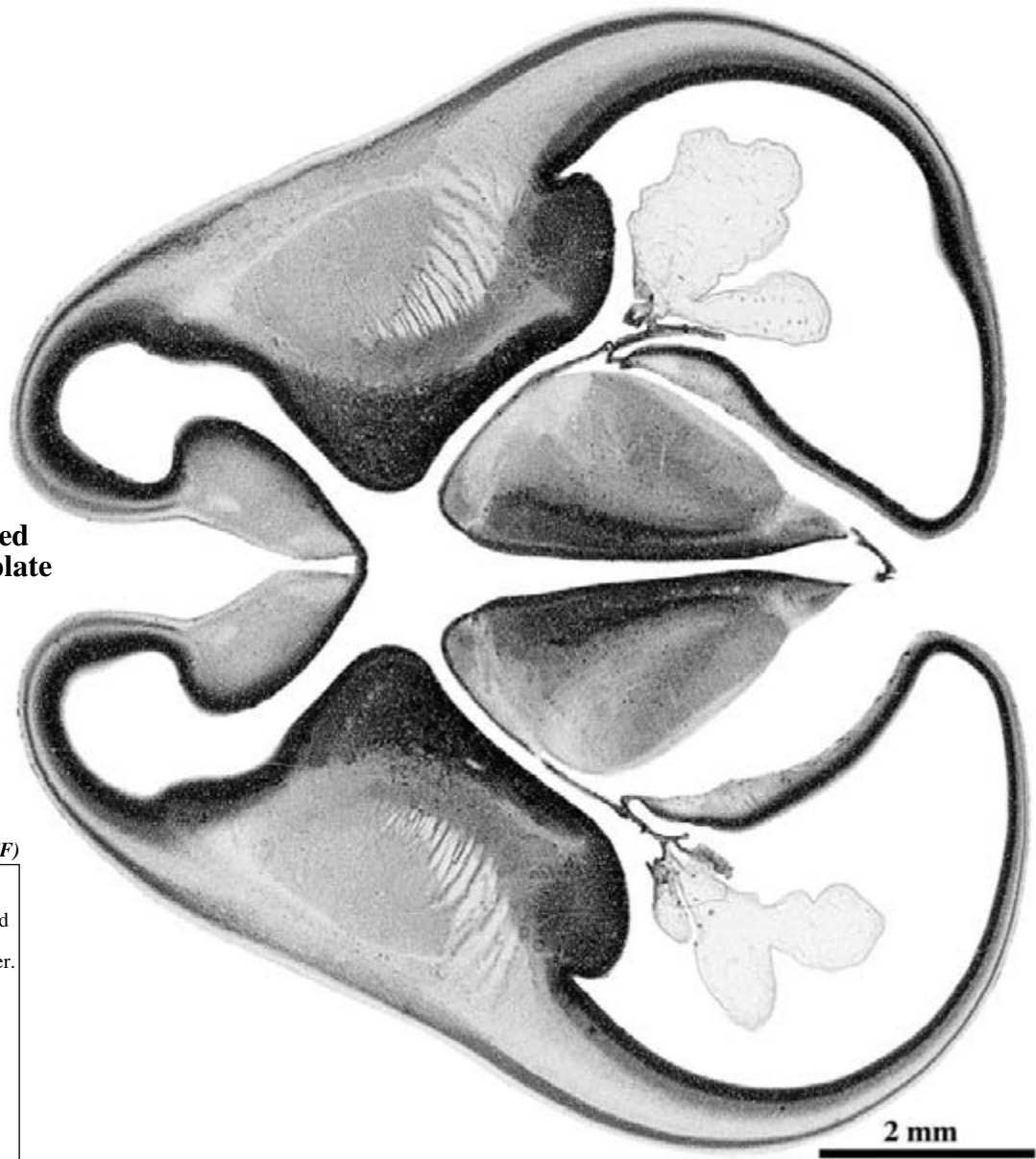


Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 86A

**GW9 Horizontal
CR 43 mm, C886
Level 4: Section 53**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



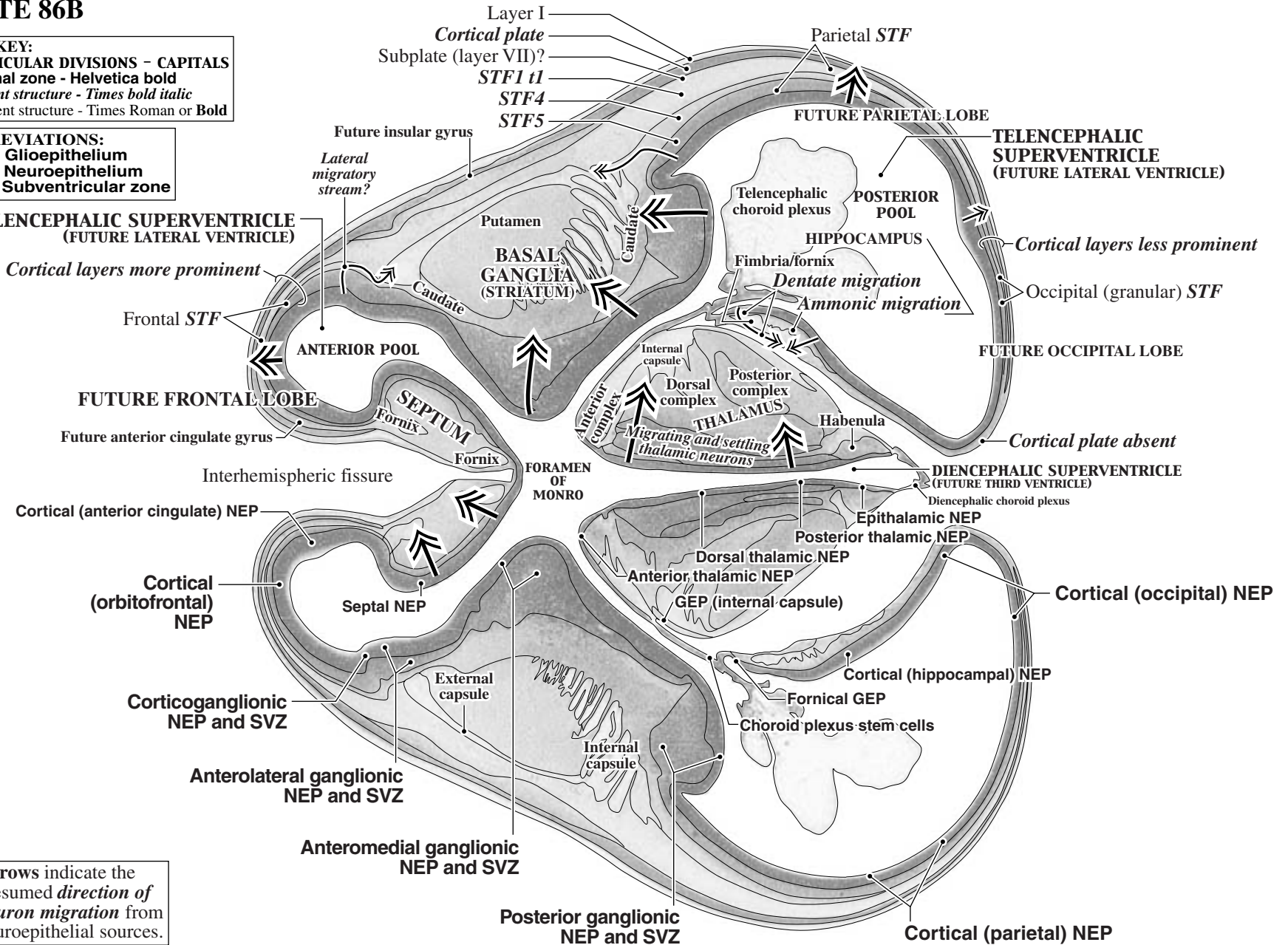
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 86B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

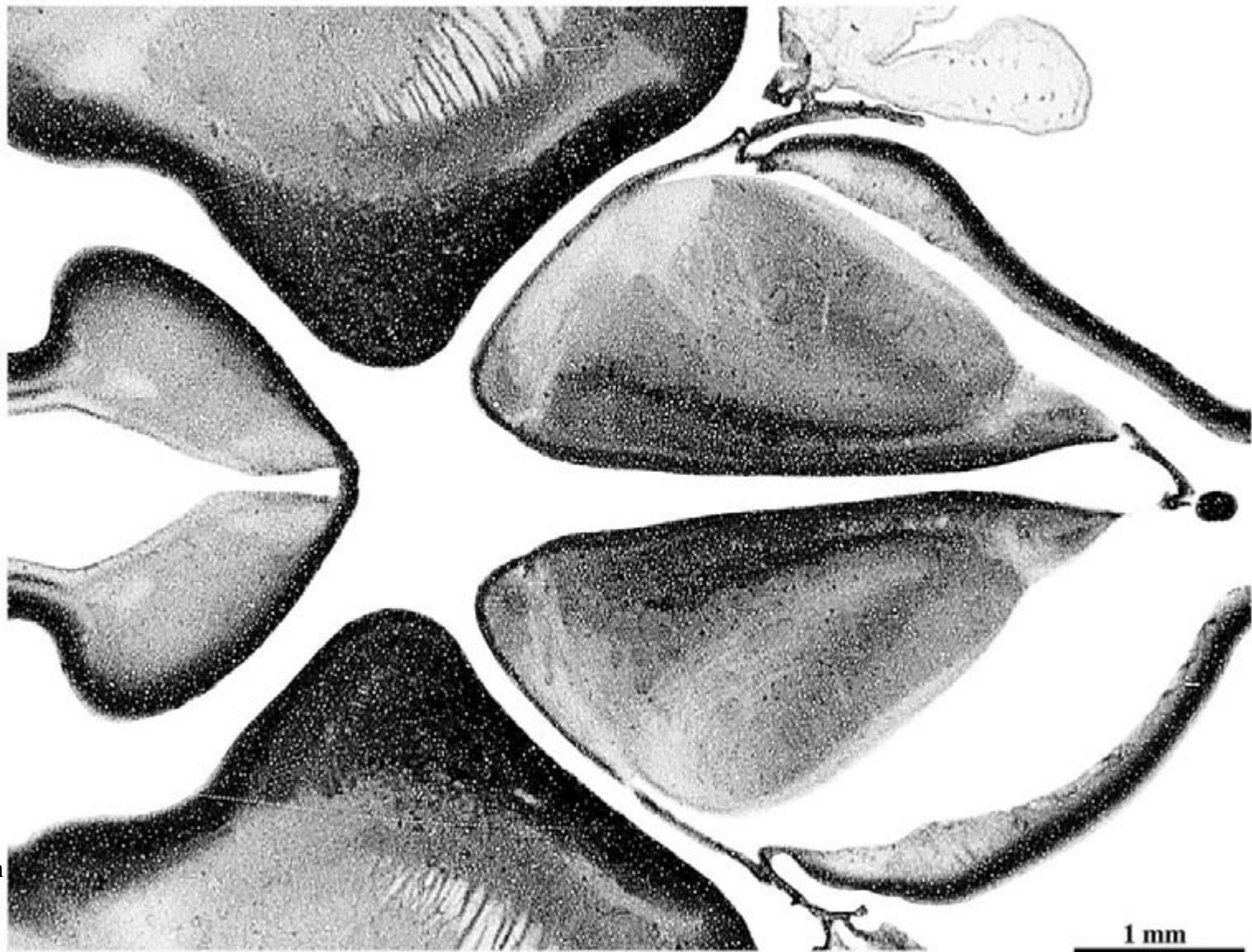
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 86C

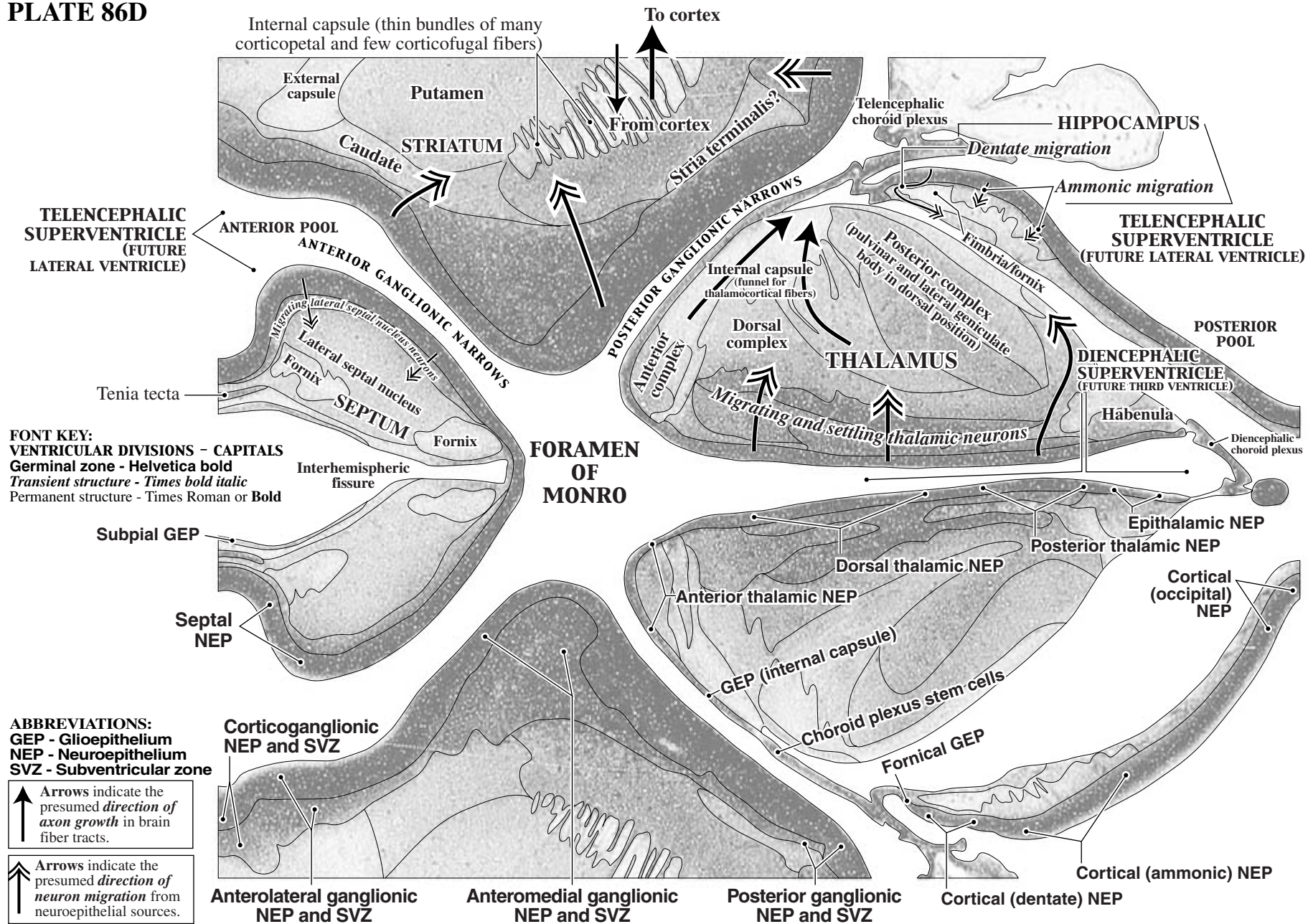
**GW9 Horizontal
CR 43 mm
C886
Level 4:
Section 53**



**See the entire section
in parts A and B of
this plate on the
preceding pages.**

1 mm

PLATE 86D



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

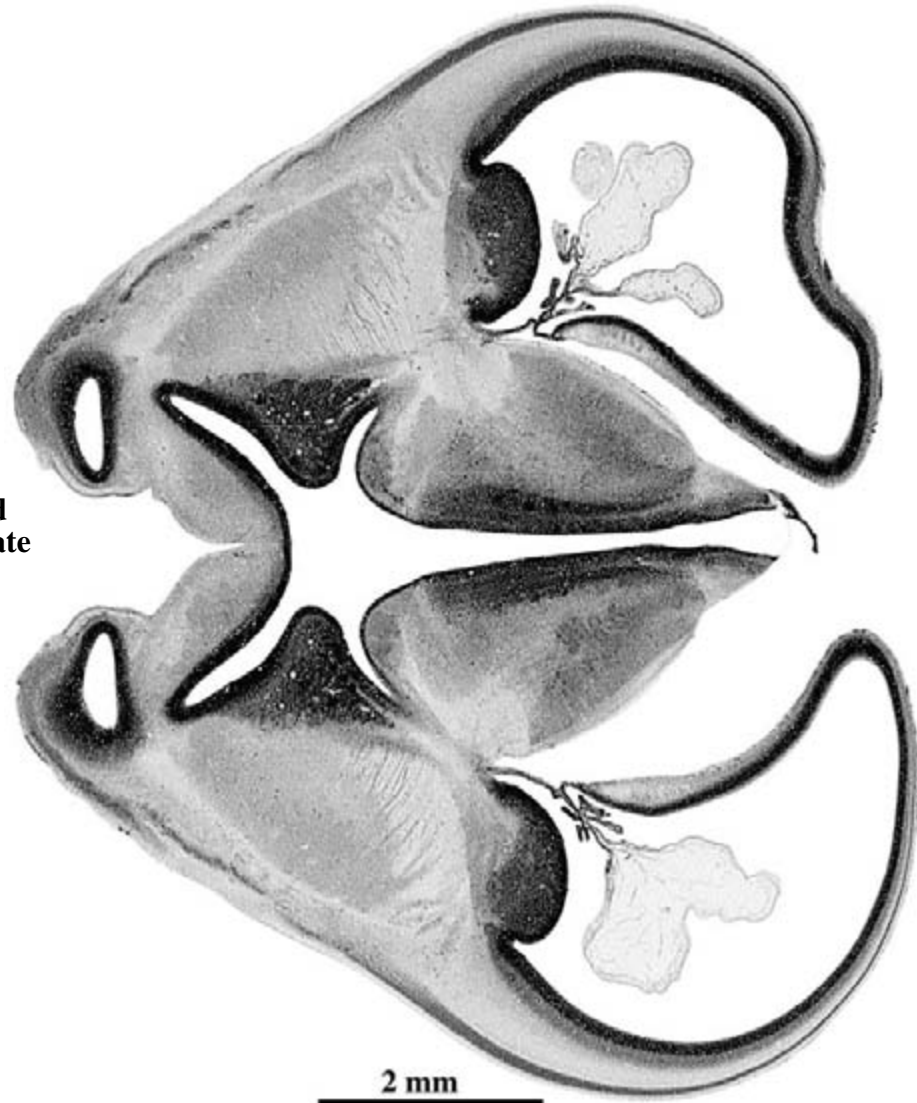
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 87A

**GW9 Horizontal
CR 43 mm, C886
Level 5: Section 57**



**See the brain core enlarged
in parts C and D of this plate
on the following pages.**

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 87B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

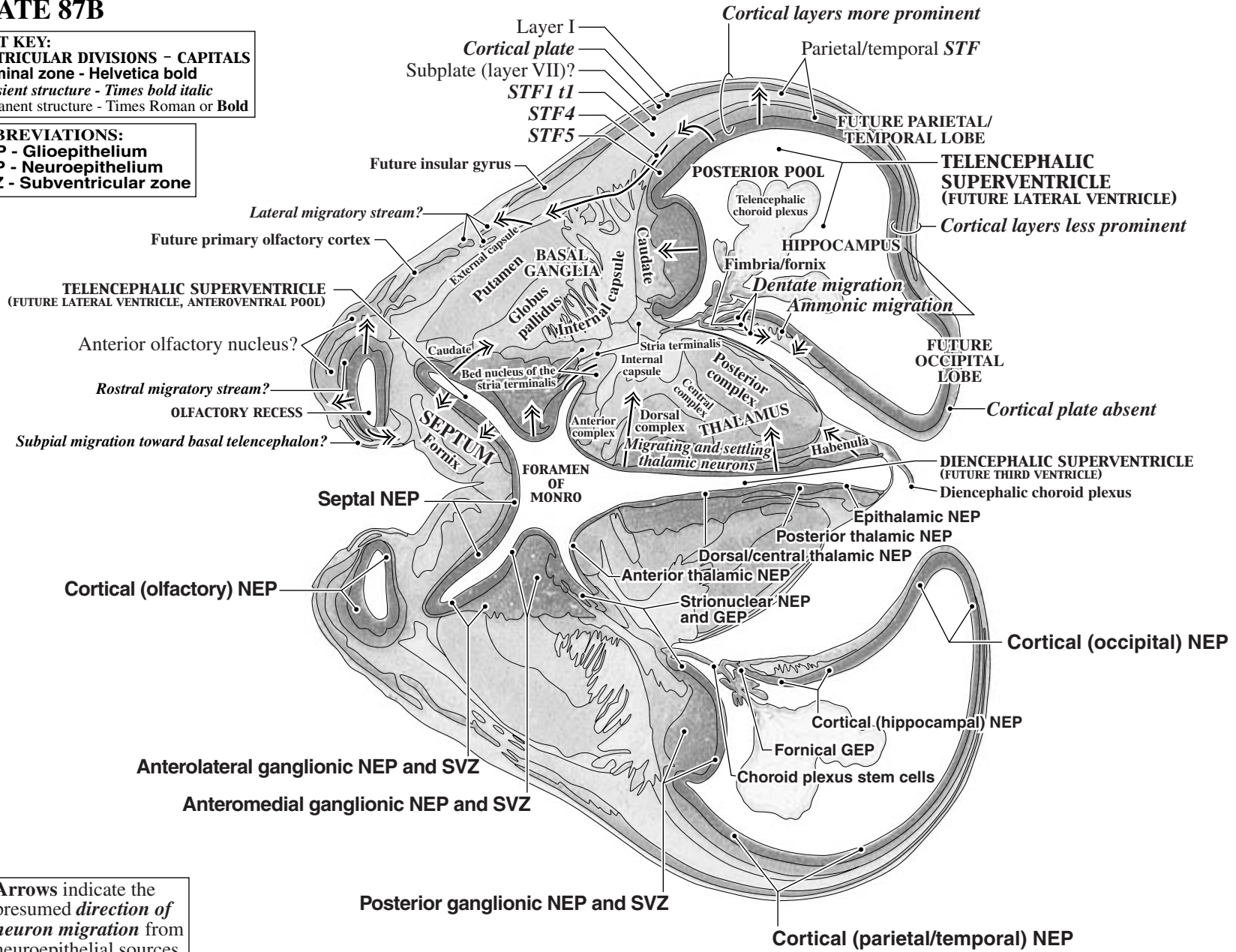
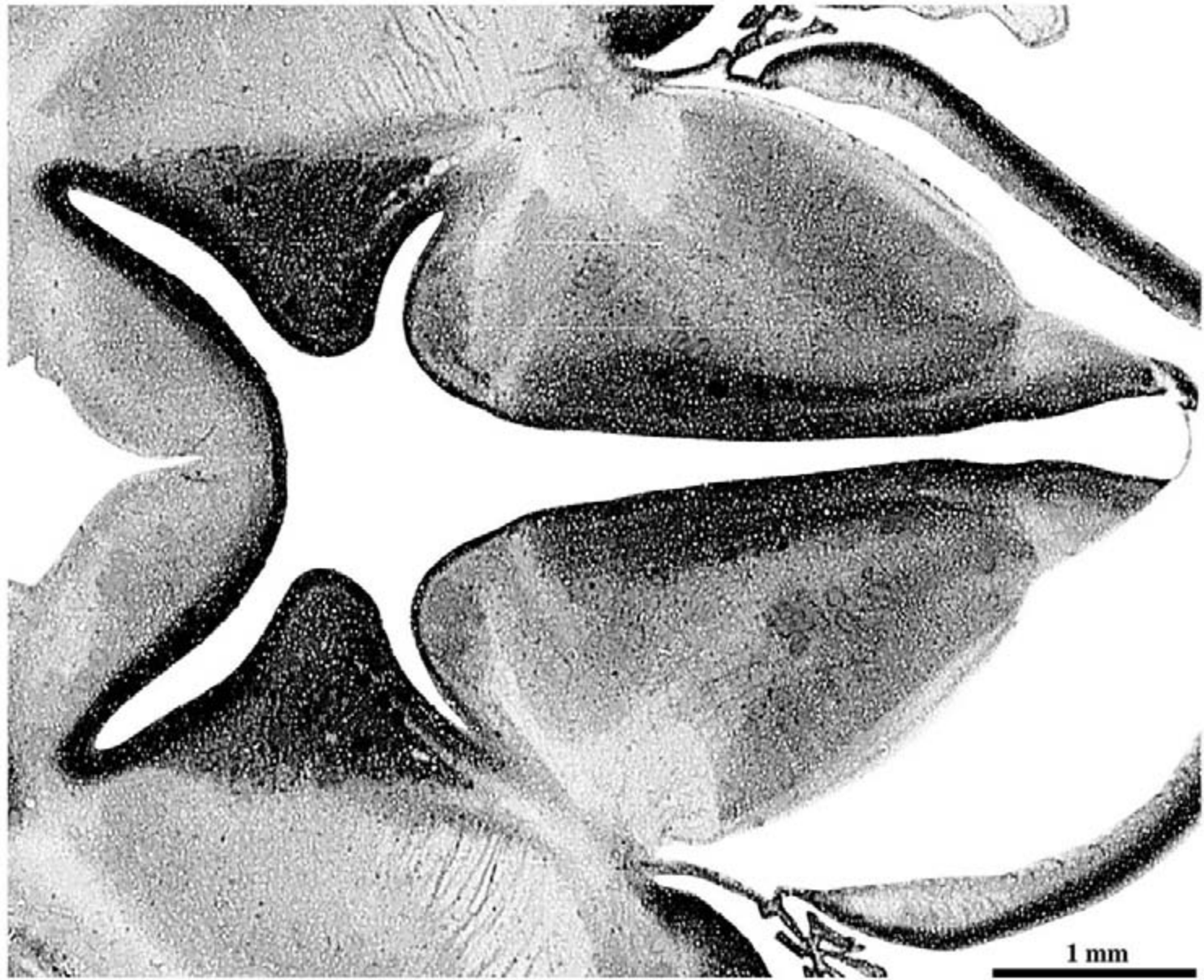


PLATE 87C

**GW9 Horizontal
CR 43 mm, C886
Level 5: Section 57**



**See the entire section
in parts A and B of
this plate on the
preceding pages.**

PLATE 87D

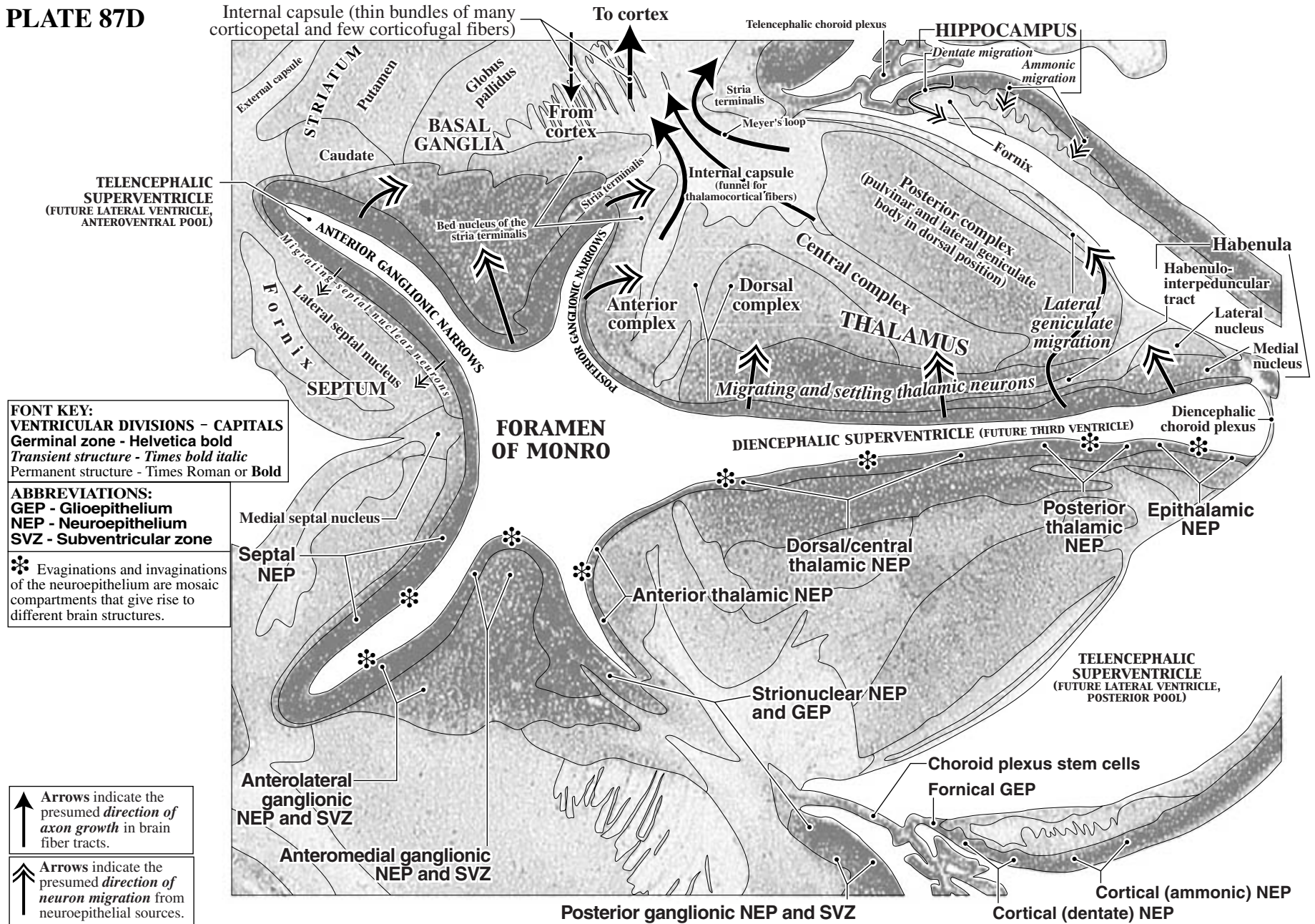


PLATE 88A

**GW9 Horizontal
CR 43 mm, C886
Level 6: Section 63**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



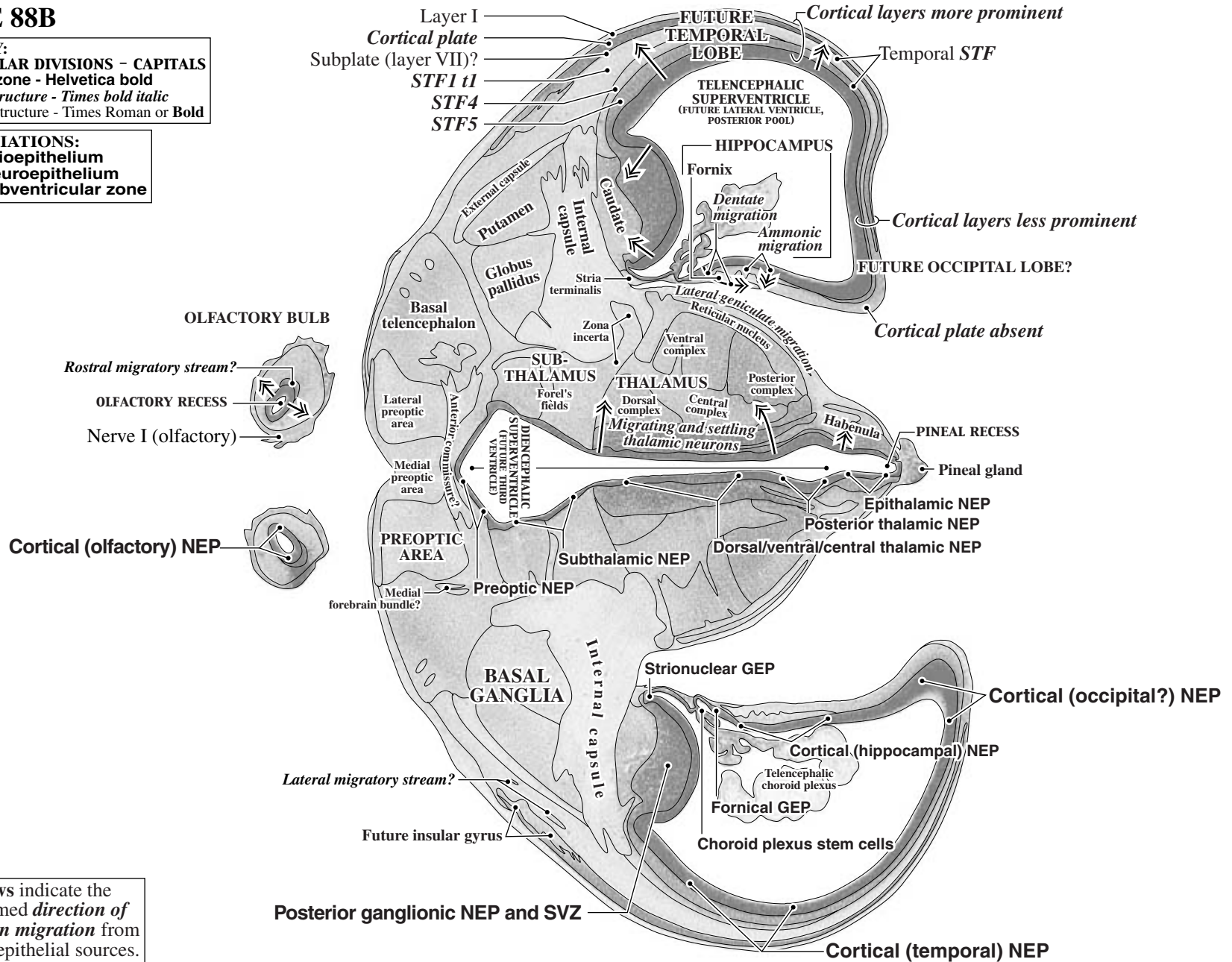
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 88B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 88C

**GW9 Horizontal
CR 43 mm, C886
Level 6: Section 63**

See the entire section
in parts A and B of
this plate on the
preceding pages.

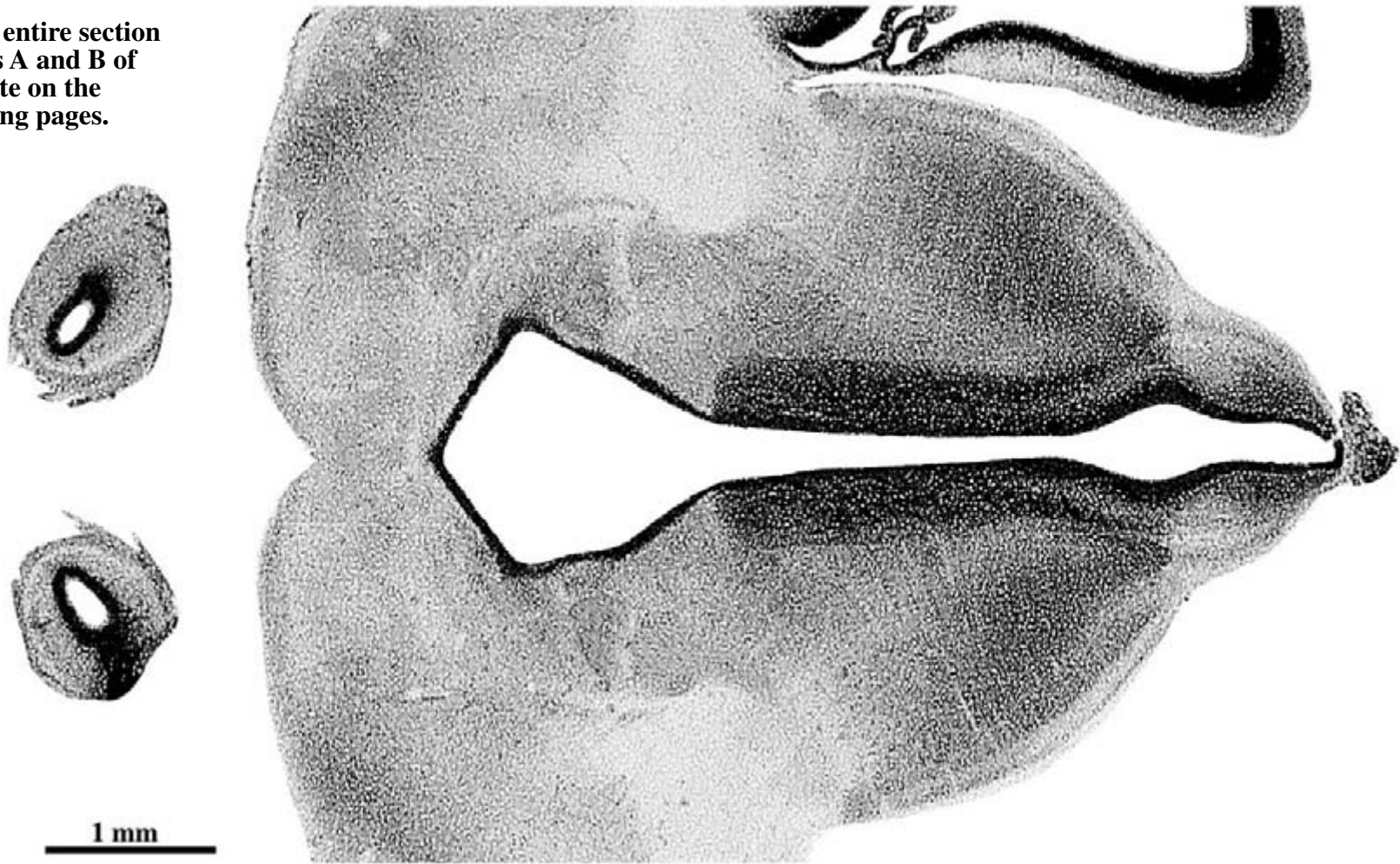


PLATE 88D

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

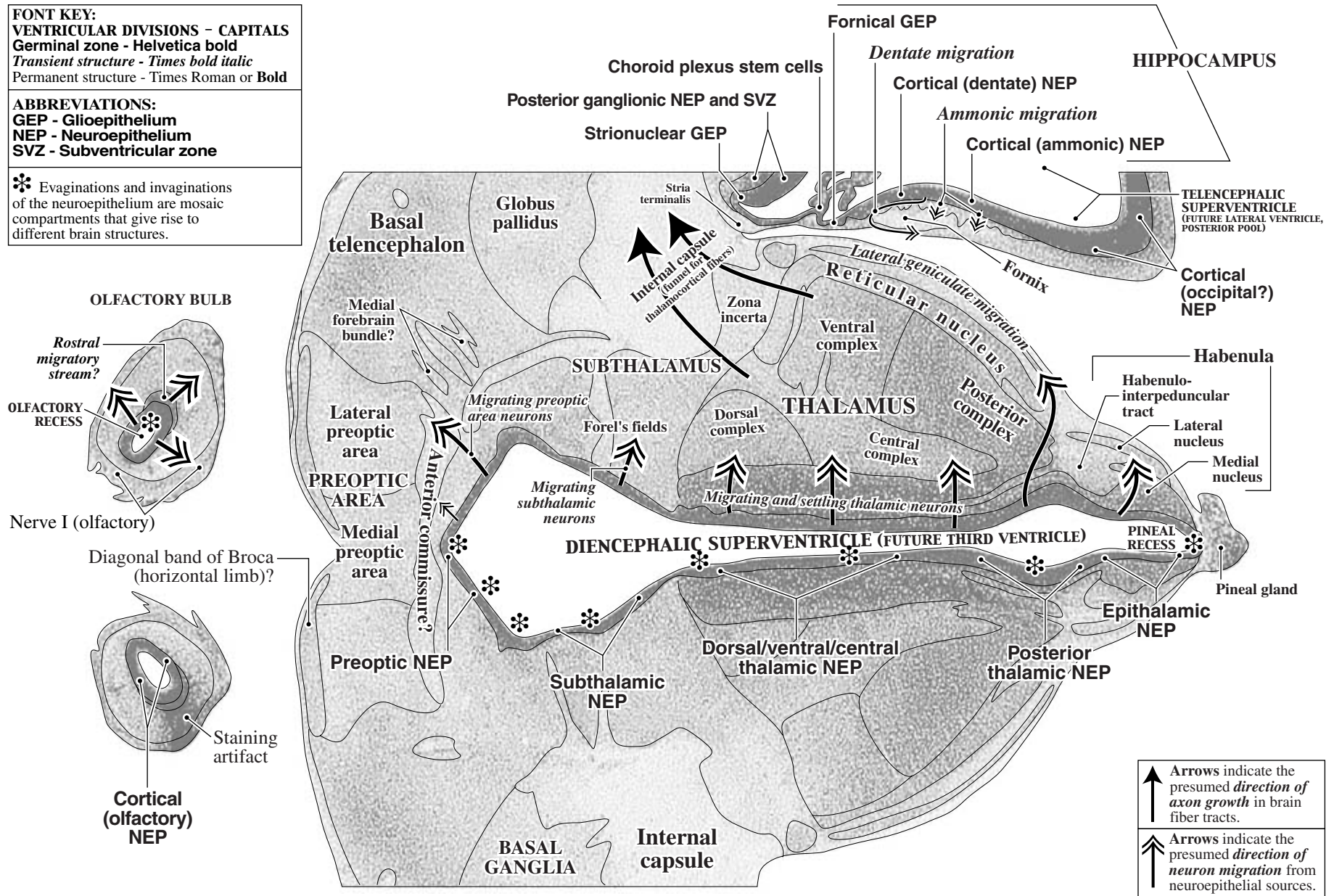
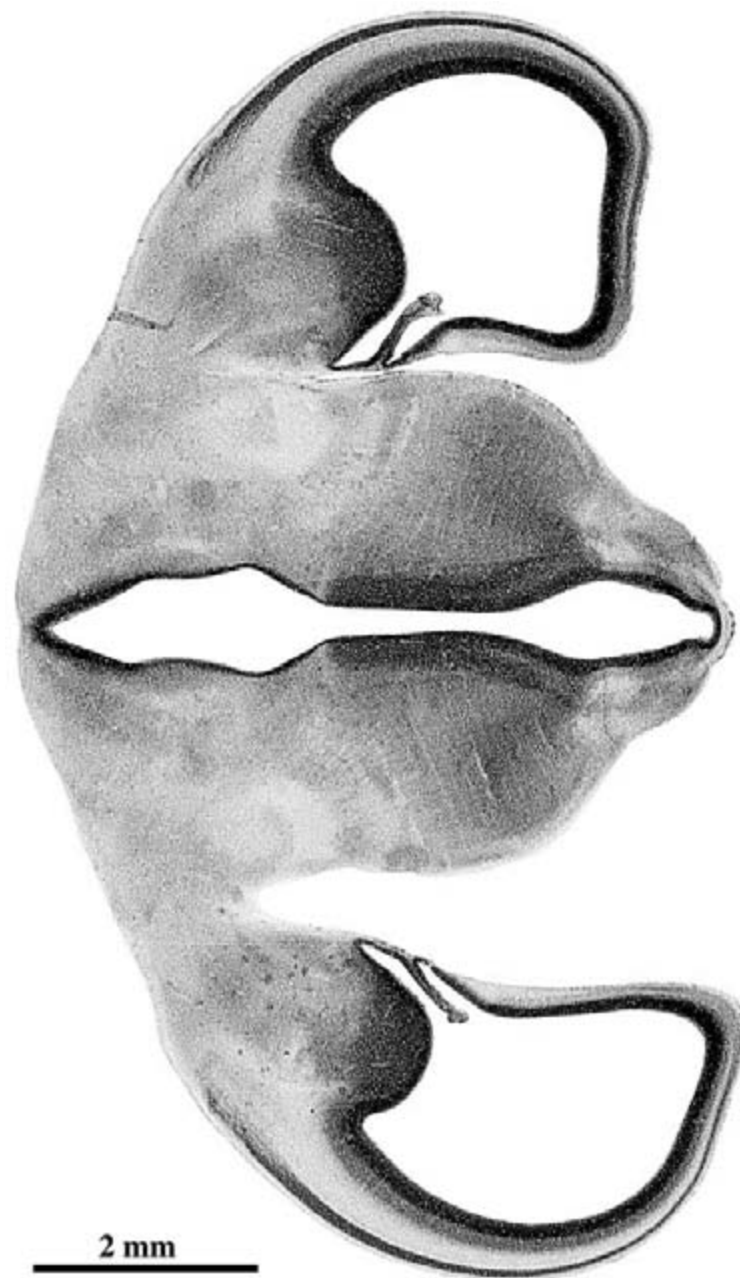


PLATE 89A

**GW9 Horizontal
CR 43 mm, C886
Level 7: Section 69**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



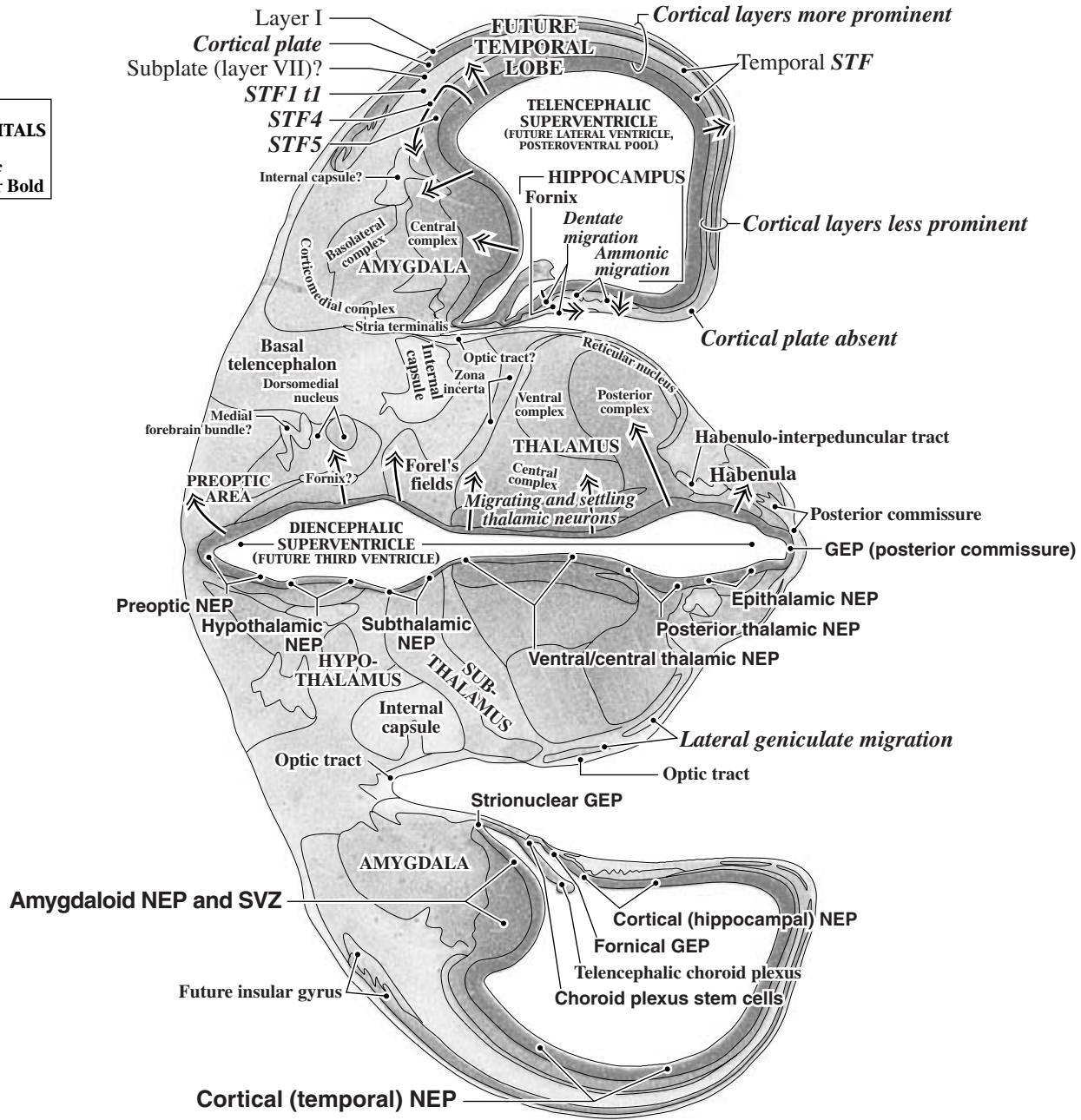
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 89B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - **Helvetica bold**
*T*ransient structure - *Times bold italic*
 Permanent structure - Times Roman or **Bold**

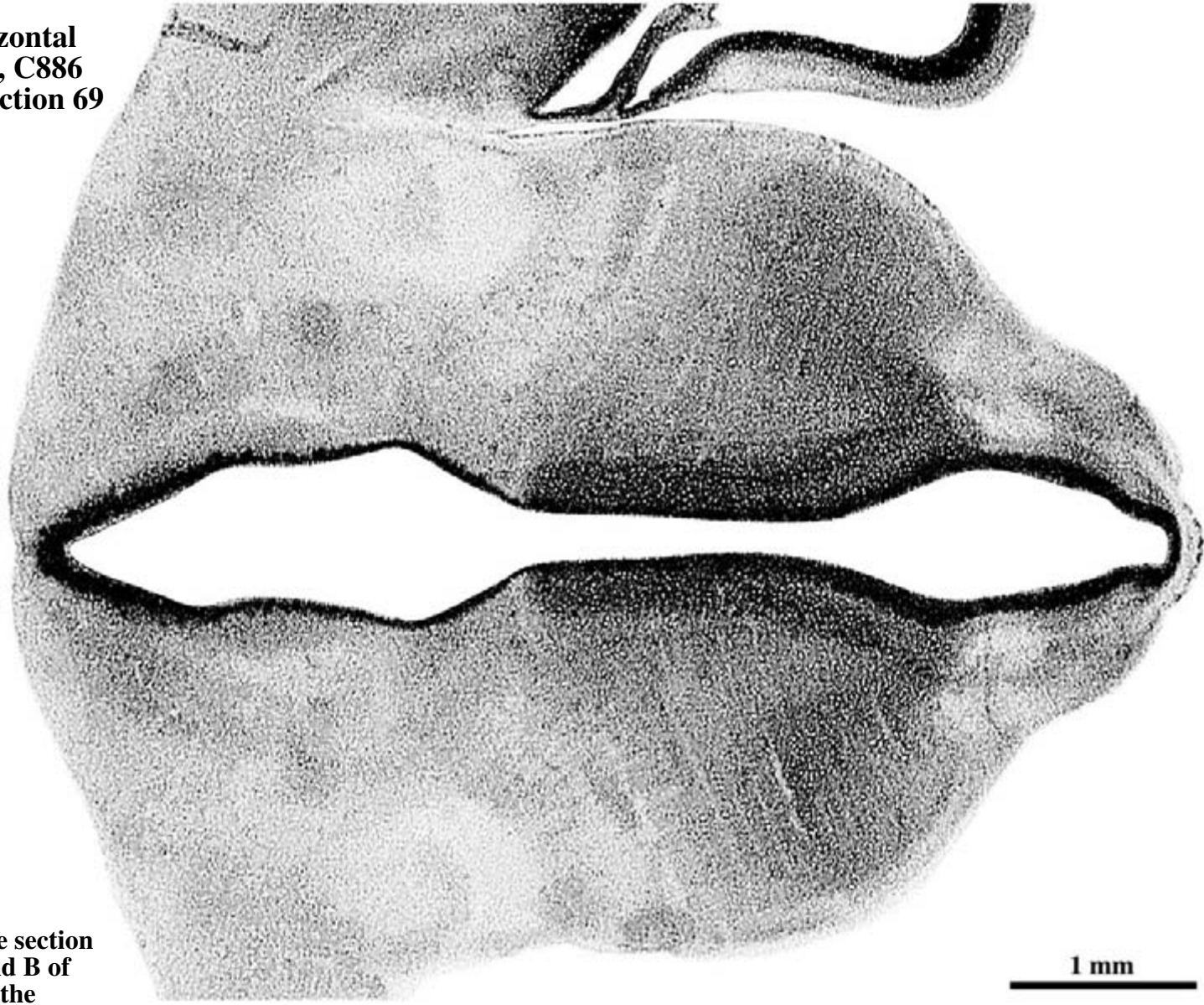
ABBREVIATIONS:
GEP - **Glioepithelium**
NEP - **Neuroepithelium**
SVZ - **Subventricular zone**



Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 89C

**GW9 Horizontal
CR 43 mm, C886
Level 7: Section 69**



**See the entire section
in parts A and B of
this plate on the
preceding pages.**

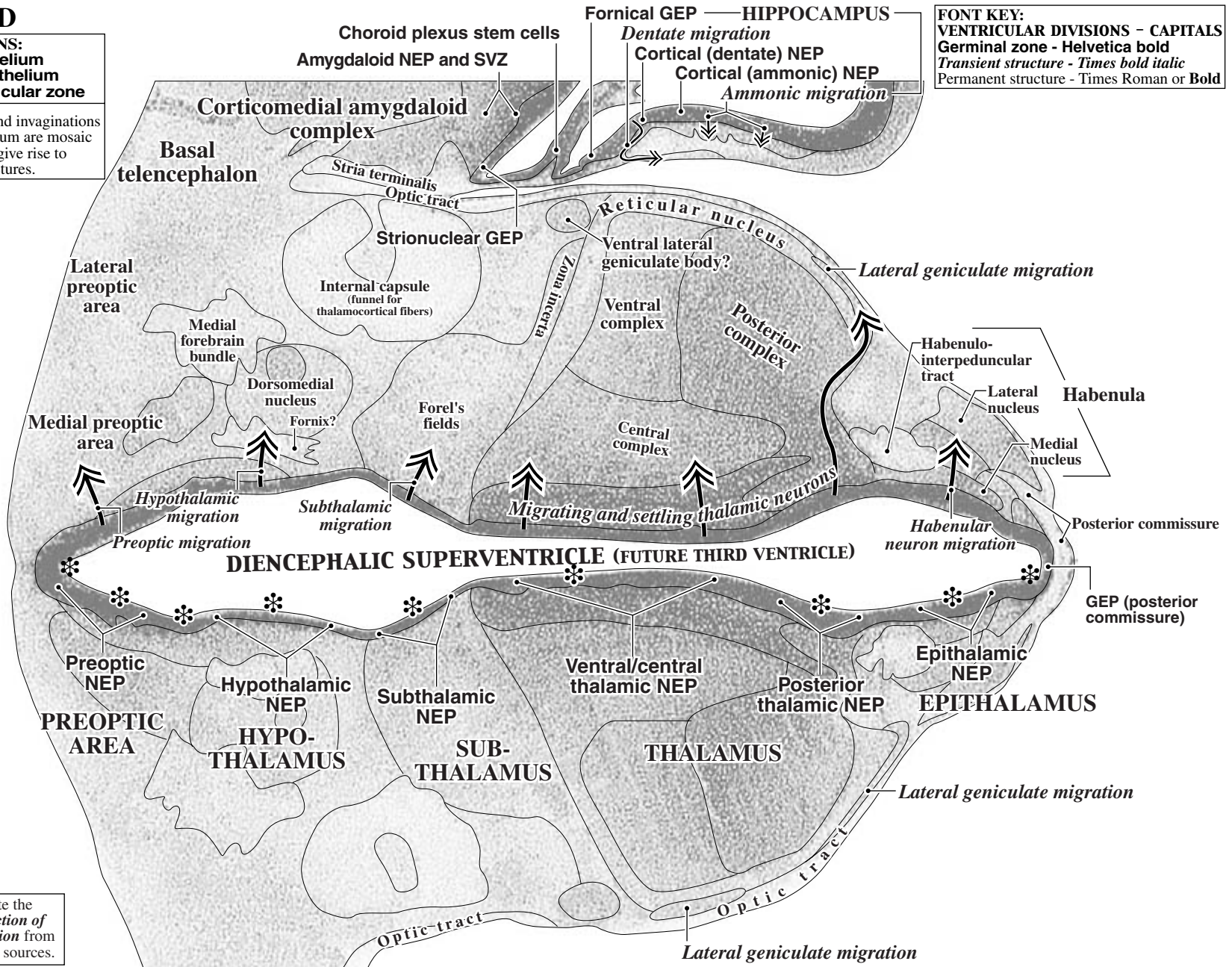
1 mm

PLATE 89D

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

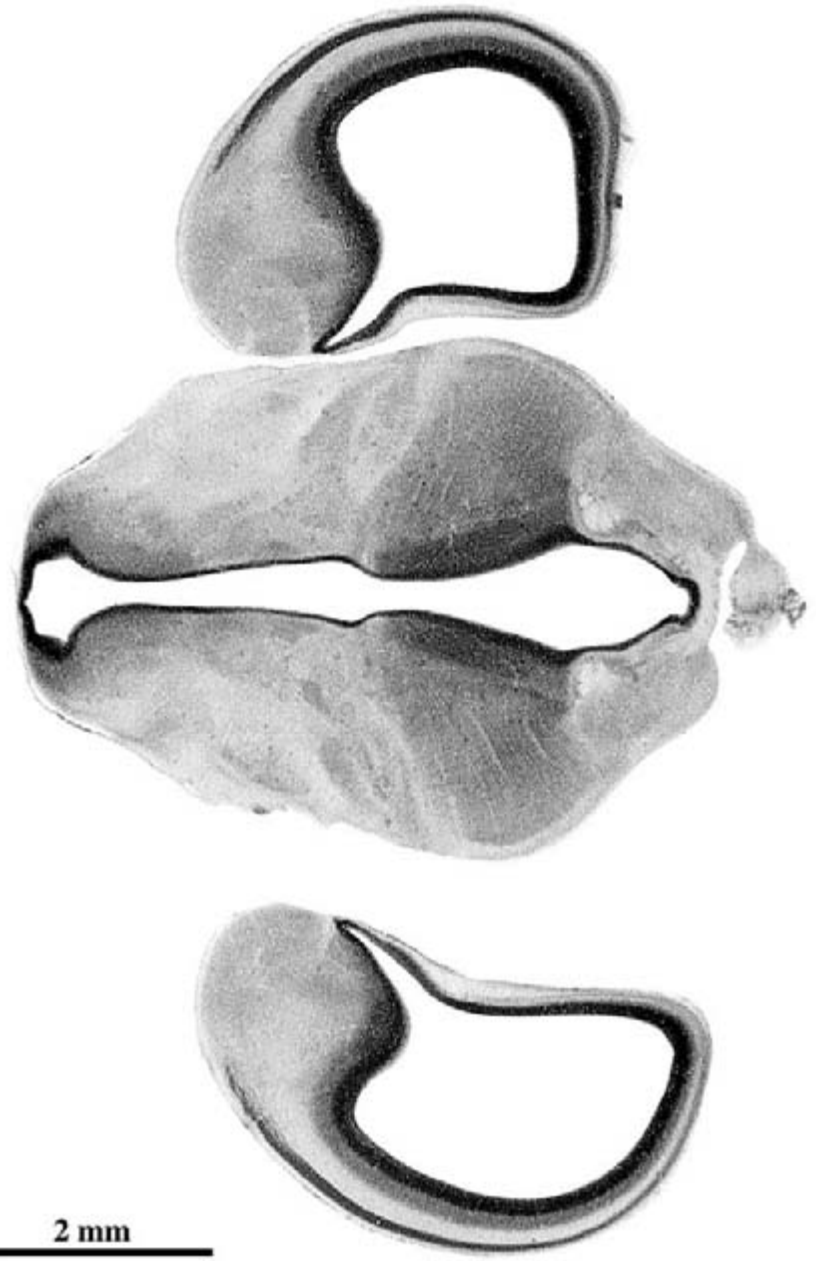


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 90A

**GW9 Horizontal
CR 43 mm, C886
Level 8: Section 75**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



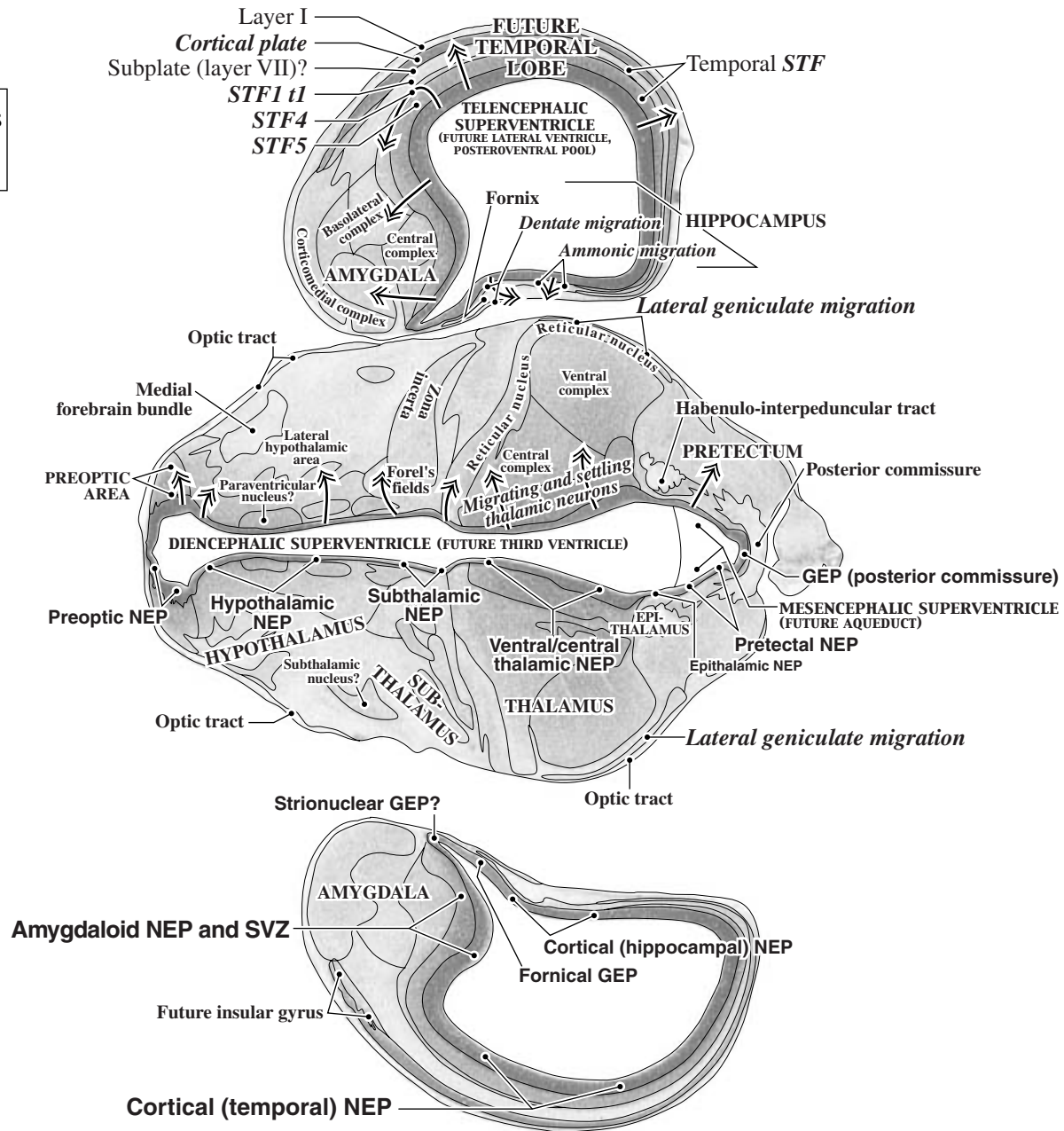
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 90B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

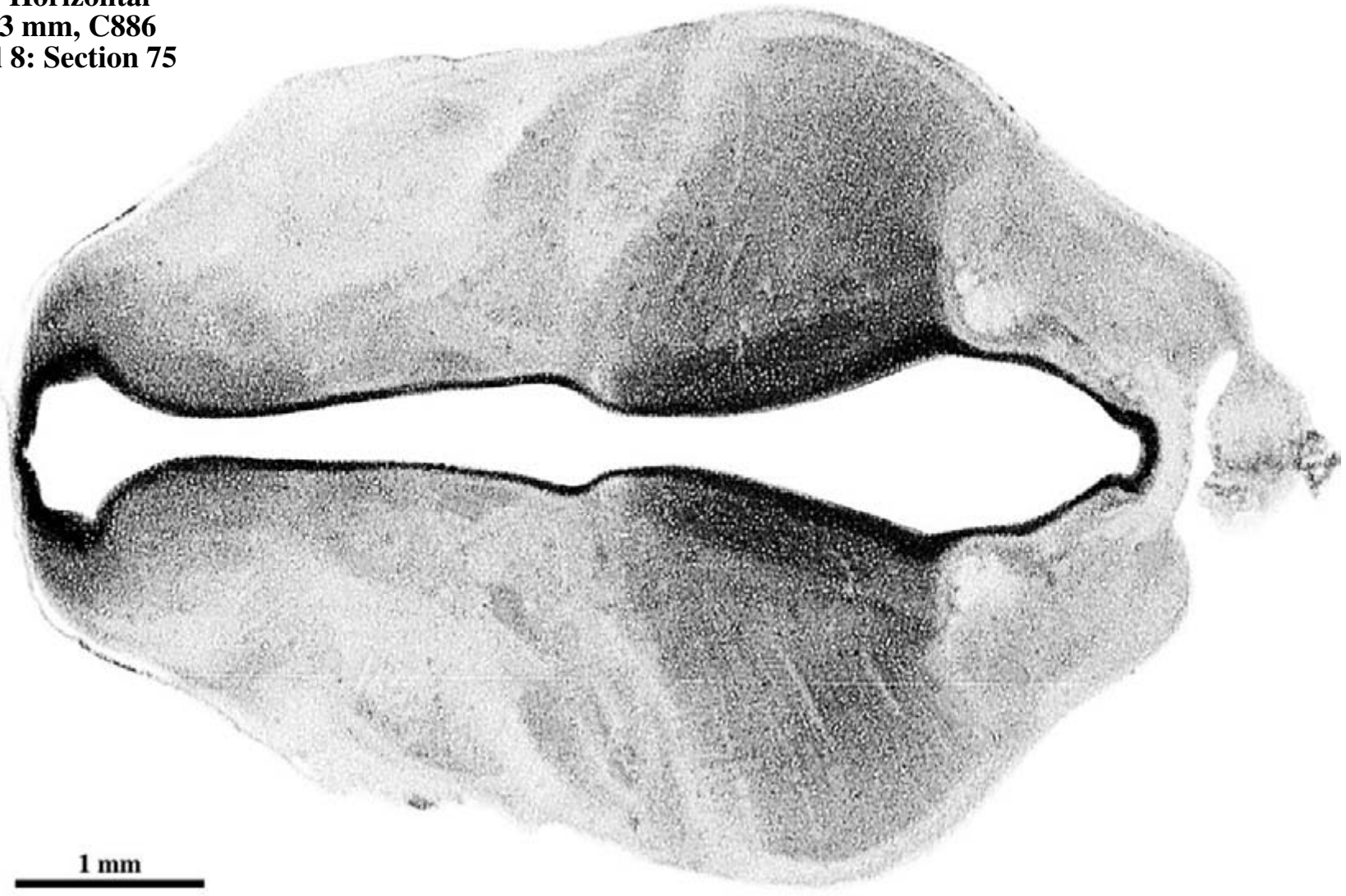
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ **Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.**

PLATE 90C

**GW9 Horizontal
CR 43 mm, C886
Level 8: Section 75**



See the entire section in parts A and B of this plate on the preceding pages.

PLATE 90D

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

✱ Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

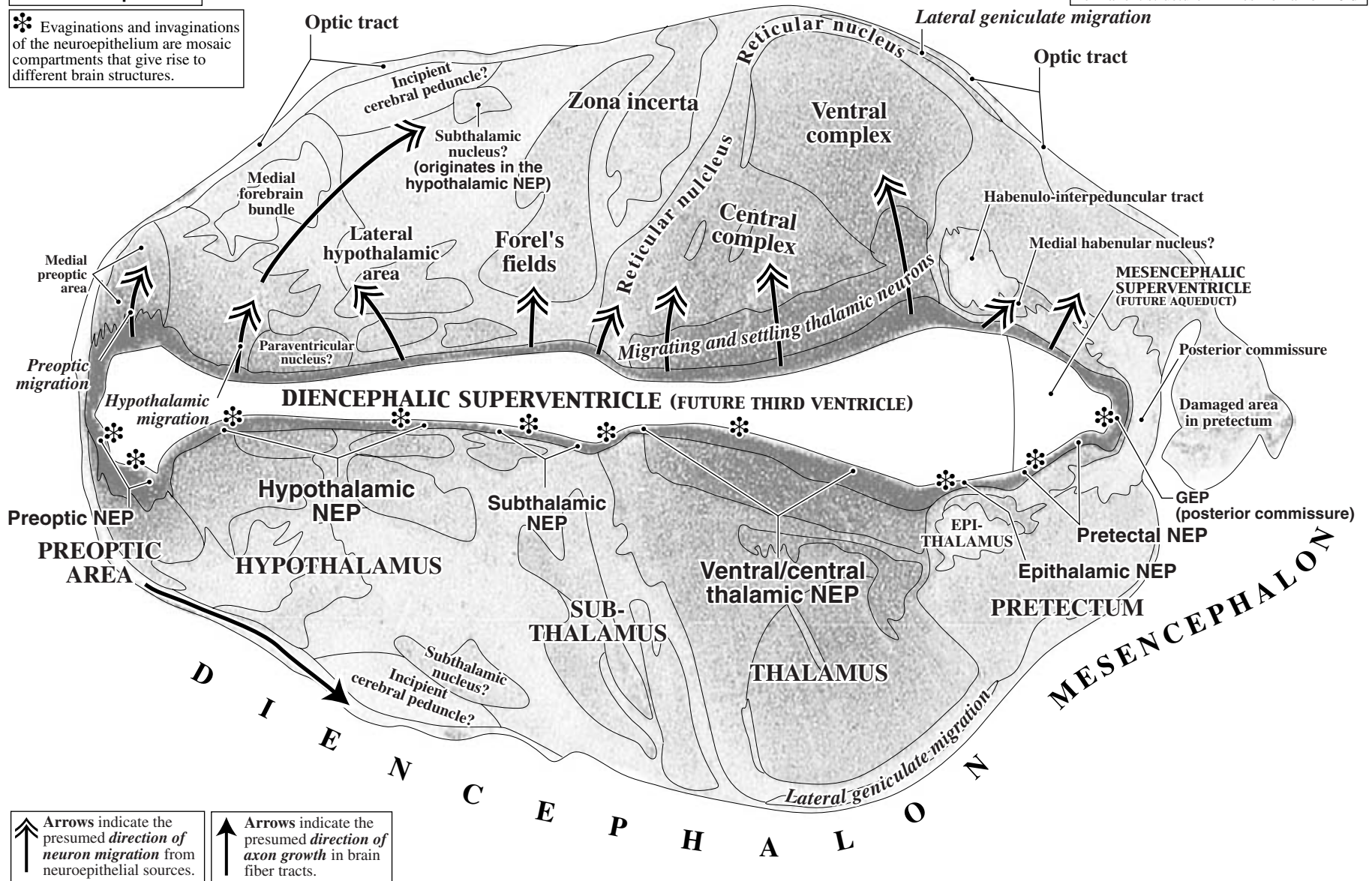
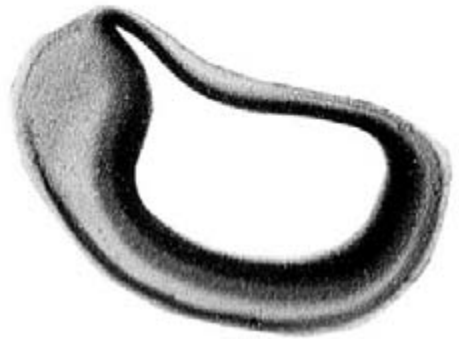
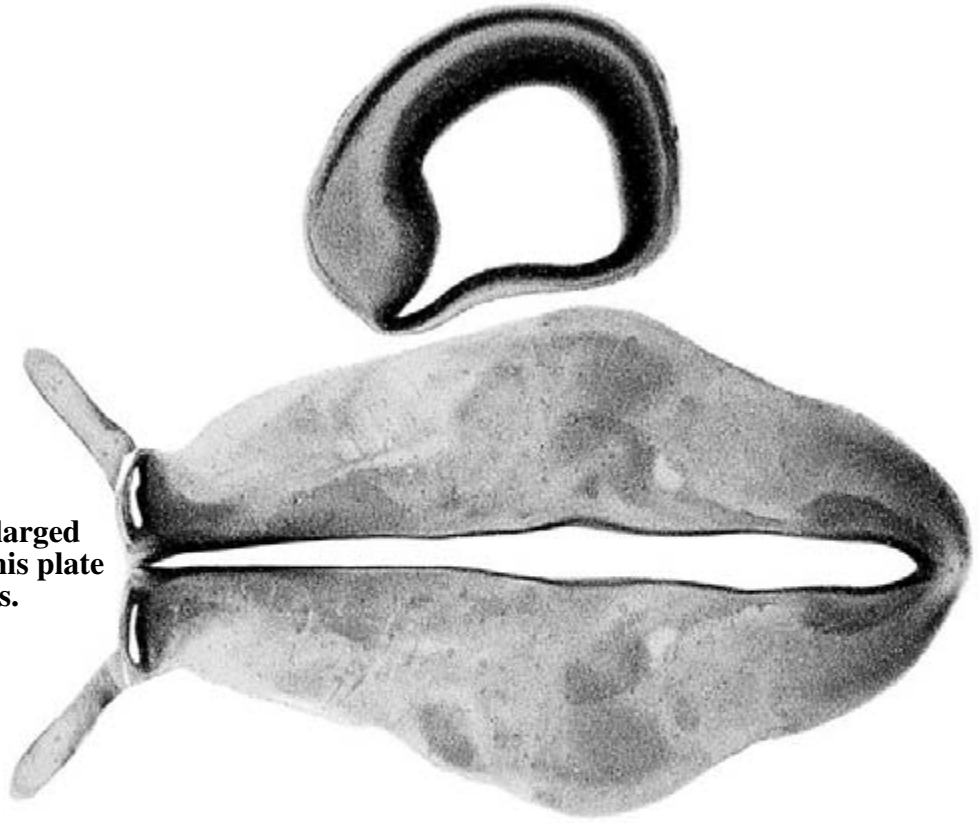


PLATE 91A

**GW9 Horizontal
CR 43 mm, C886
Level 9: Section 81**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



2 mm

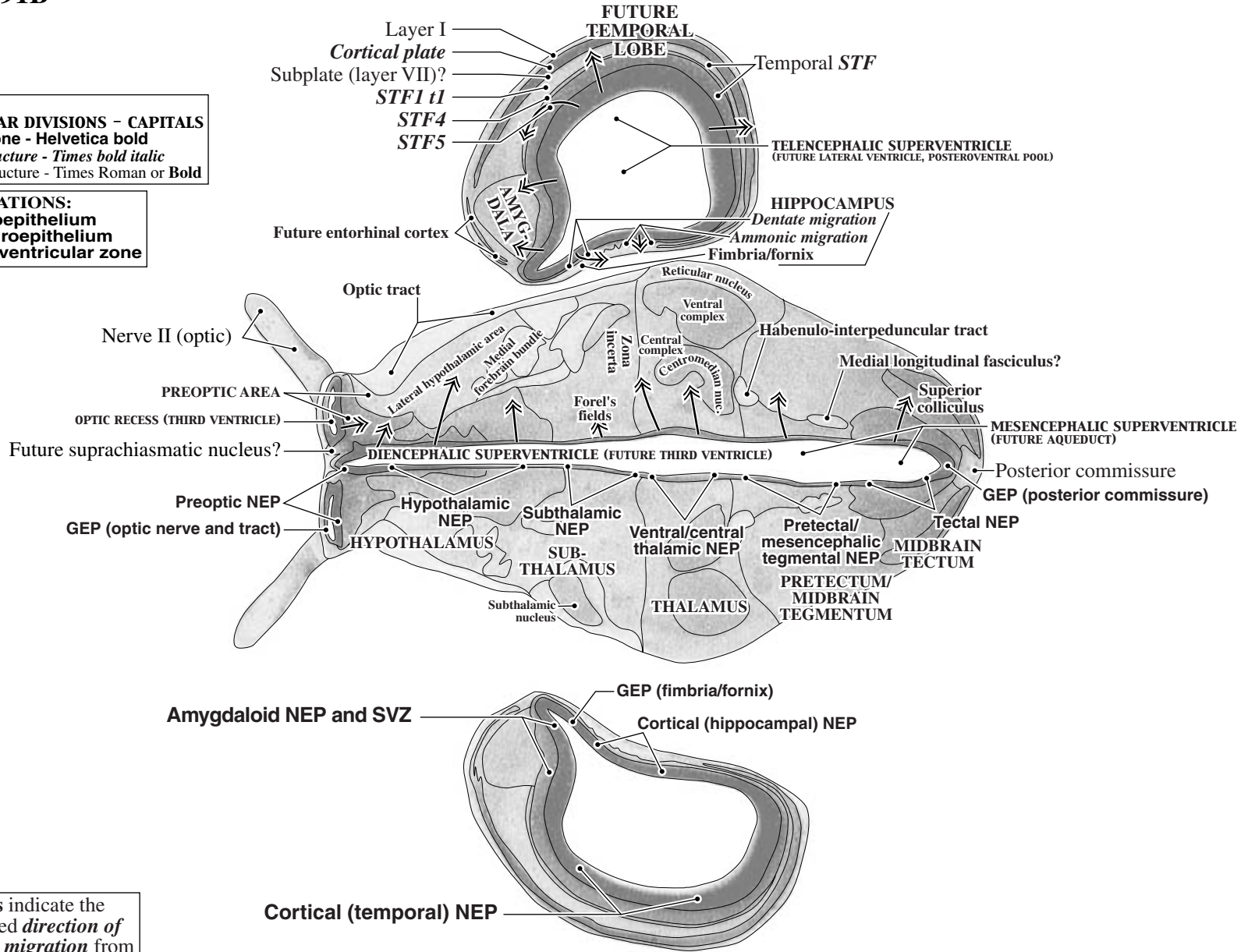
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 91B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

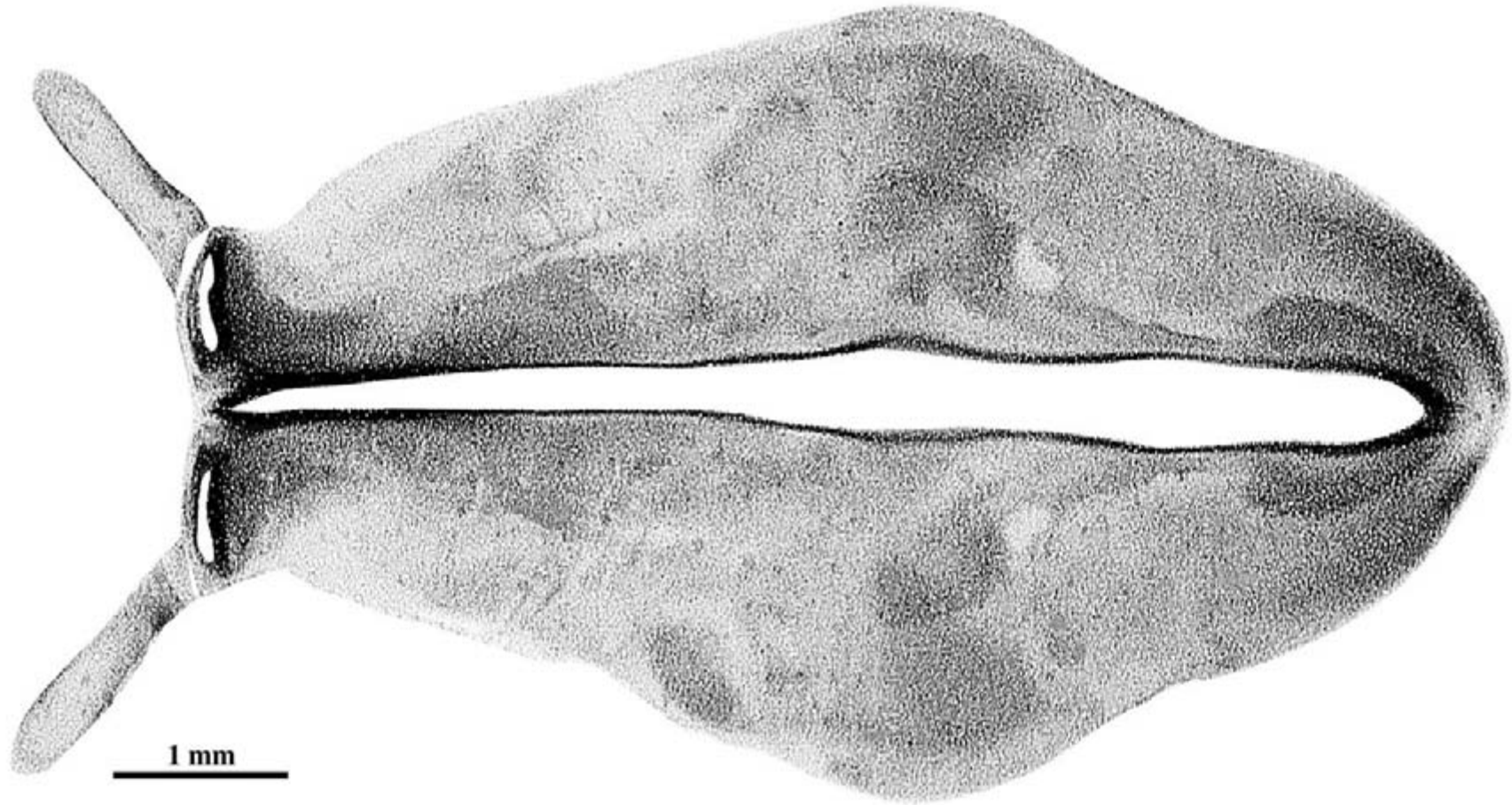
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 91C

**GW9 Horizontal
CR 43 mm, C886
Level 9: Section 81**



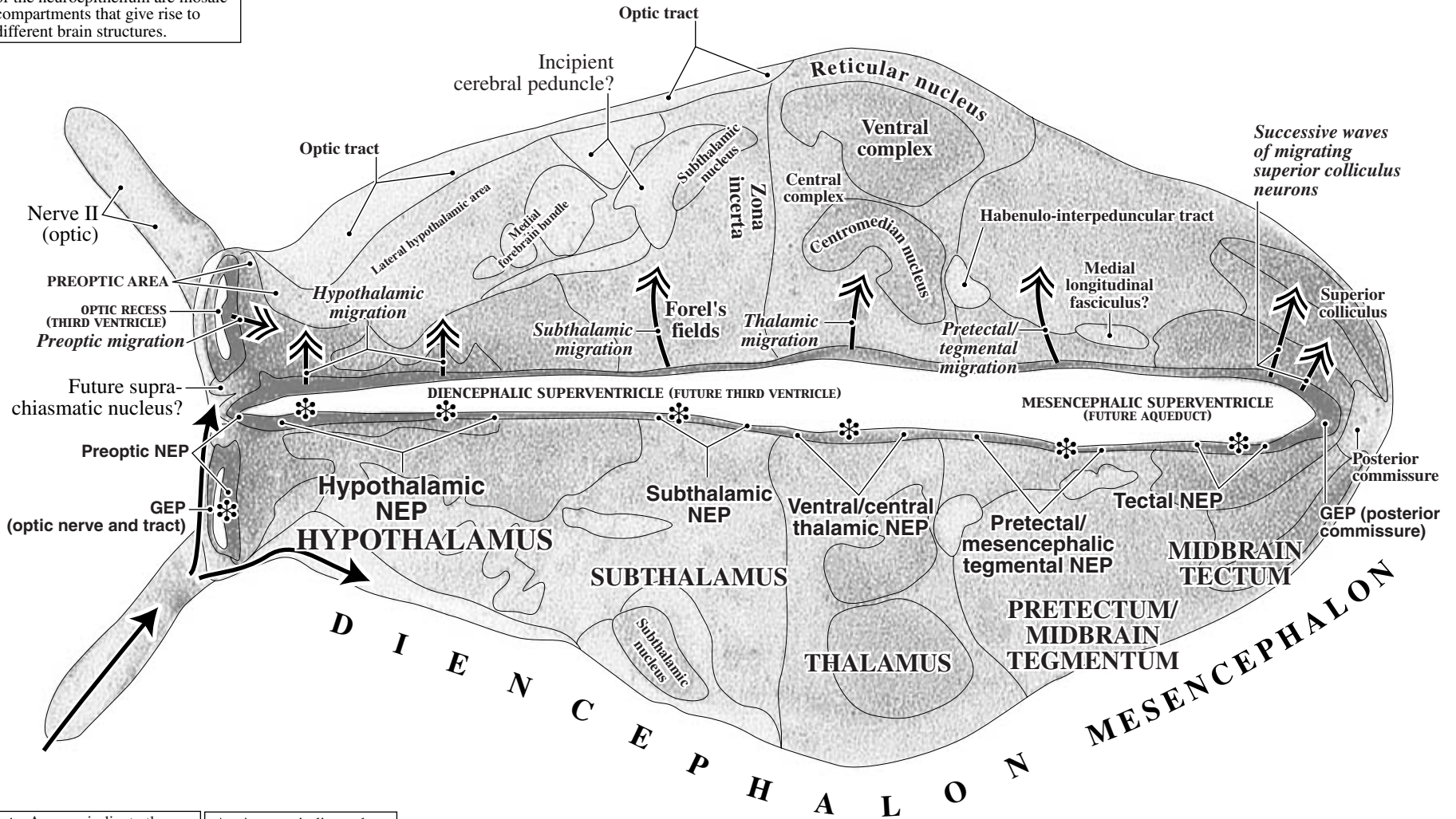
See the entire section in parts A and B of this plate on the preceding pages.

PLATE 91D

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
GERMINAL ZONE - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold



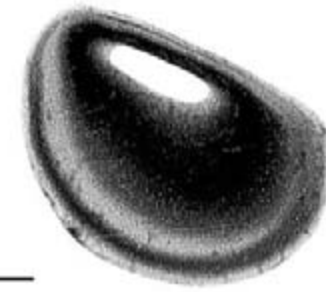
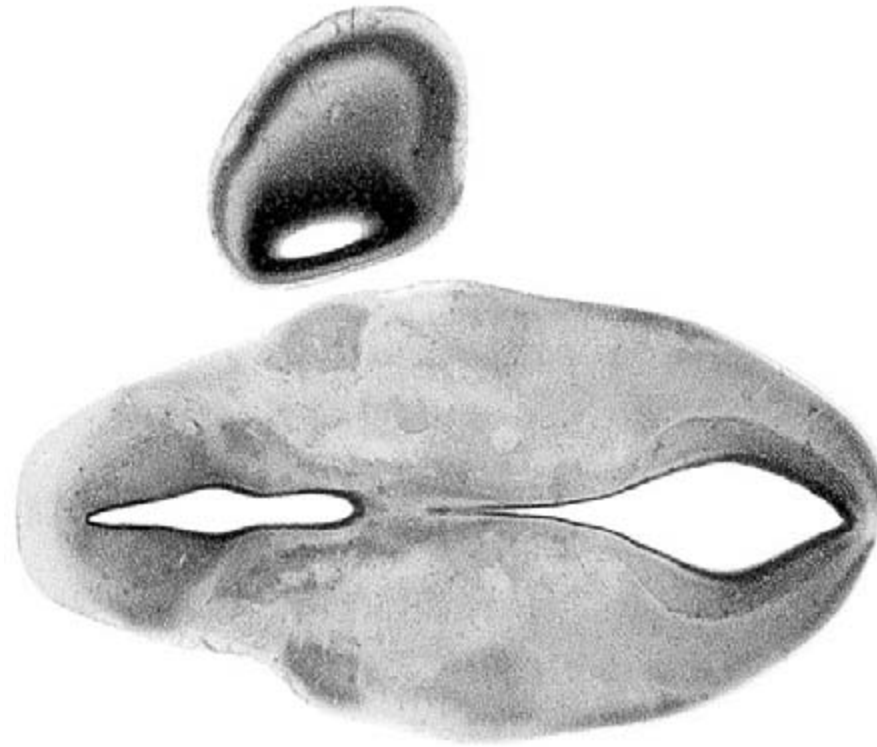
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 92A

**GW9 Horizontal
CR 43 mm, C886
Level 10: Section 86**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**



2 mm

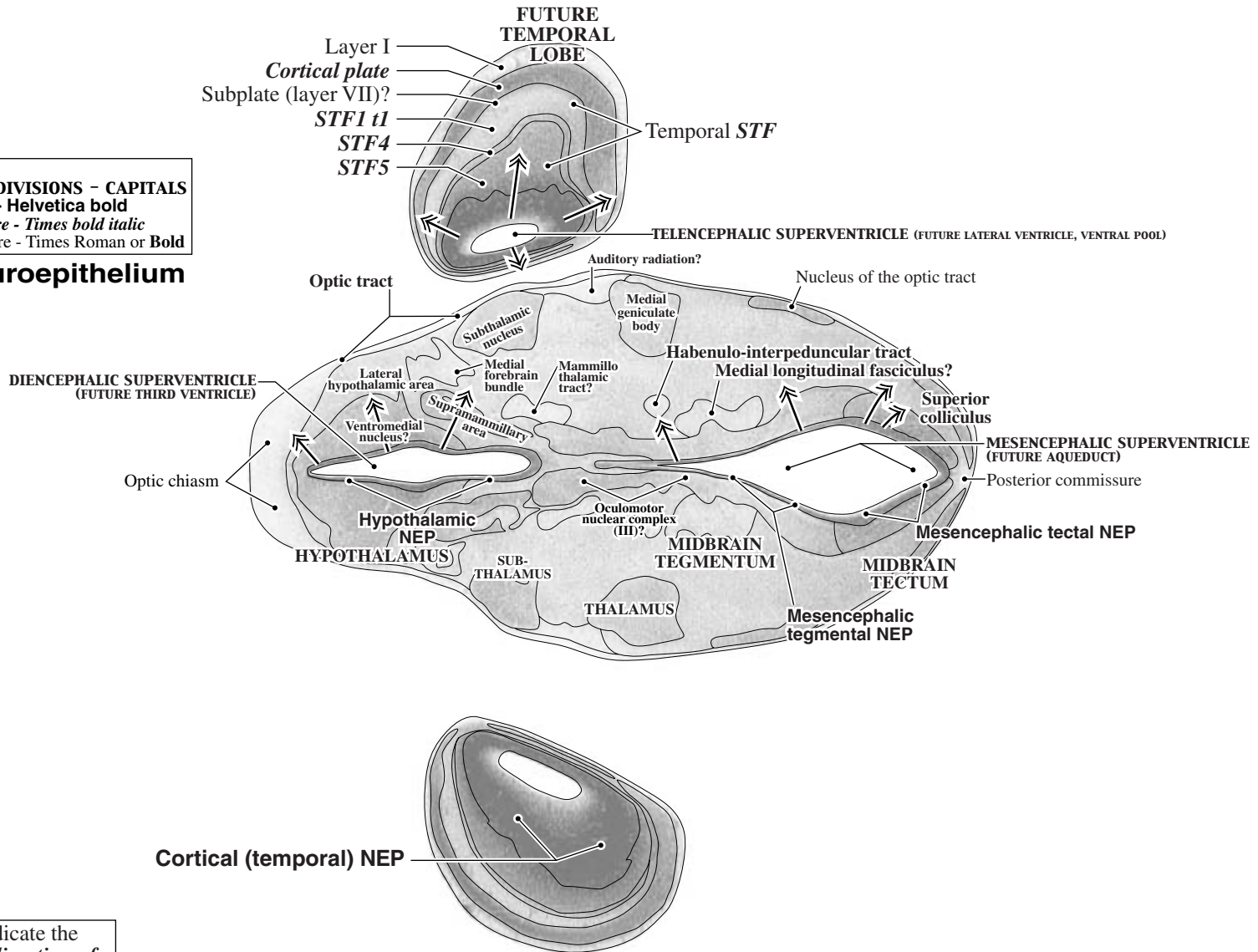
LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 92B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

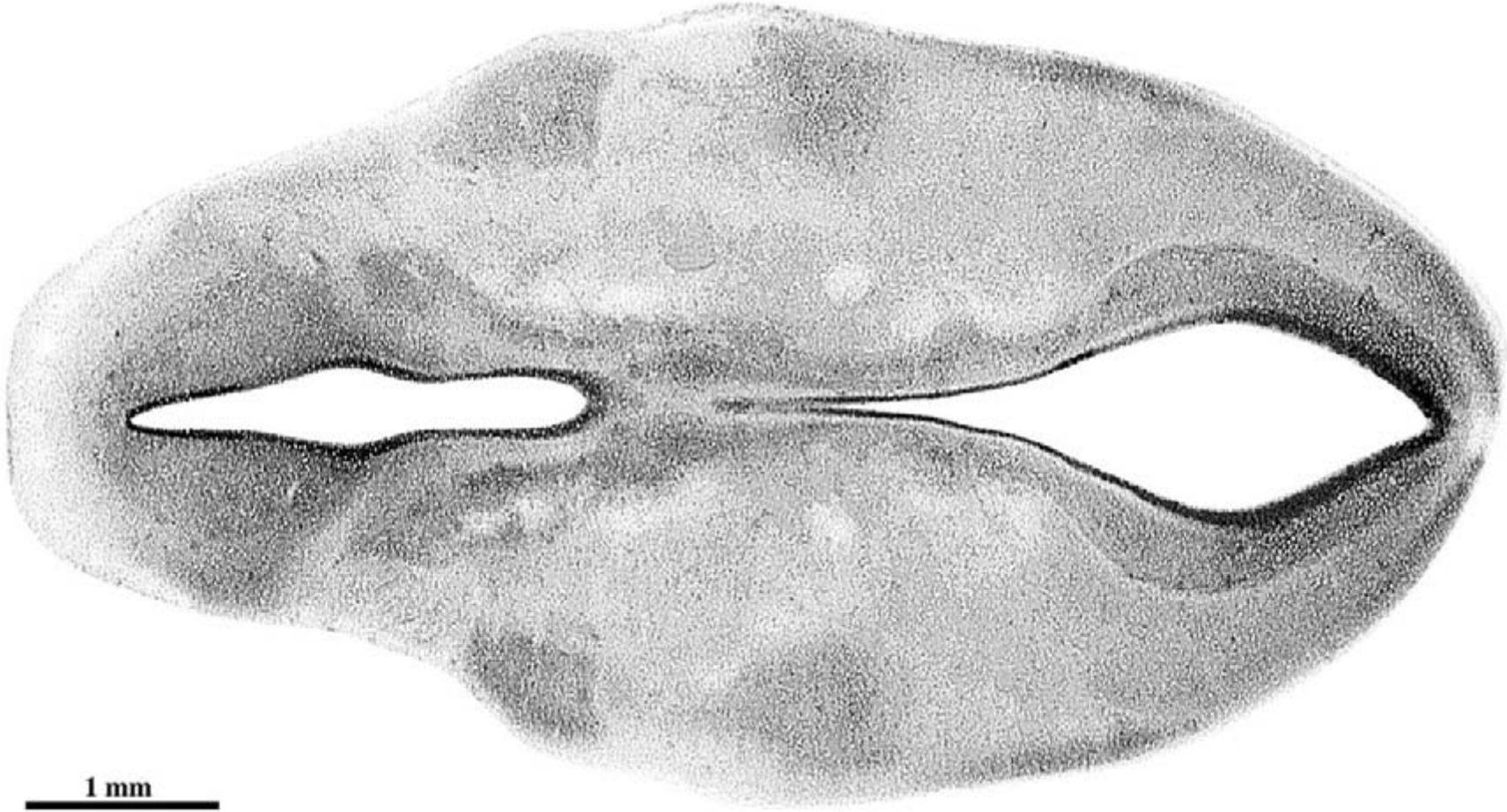
NEP - Neuroepithelium



Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 92C

**GW9 Horizontal
CR 43 mm, C886
Level 10: Section 86**



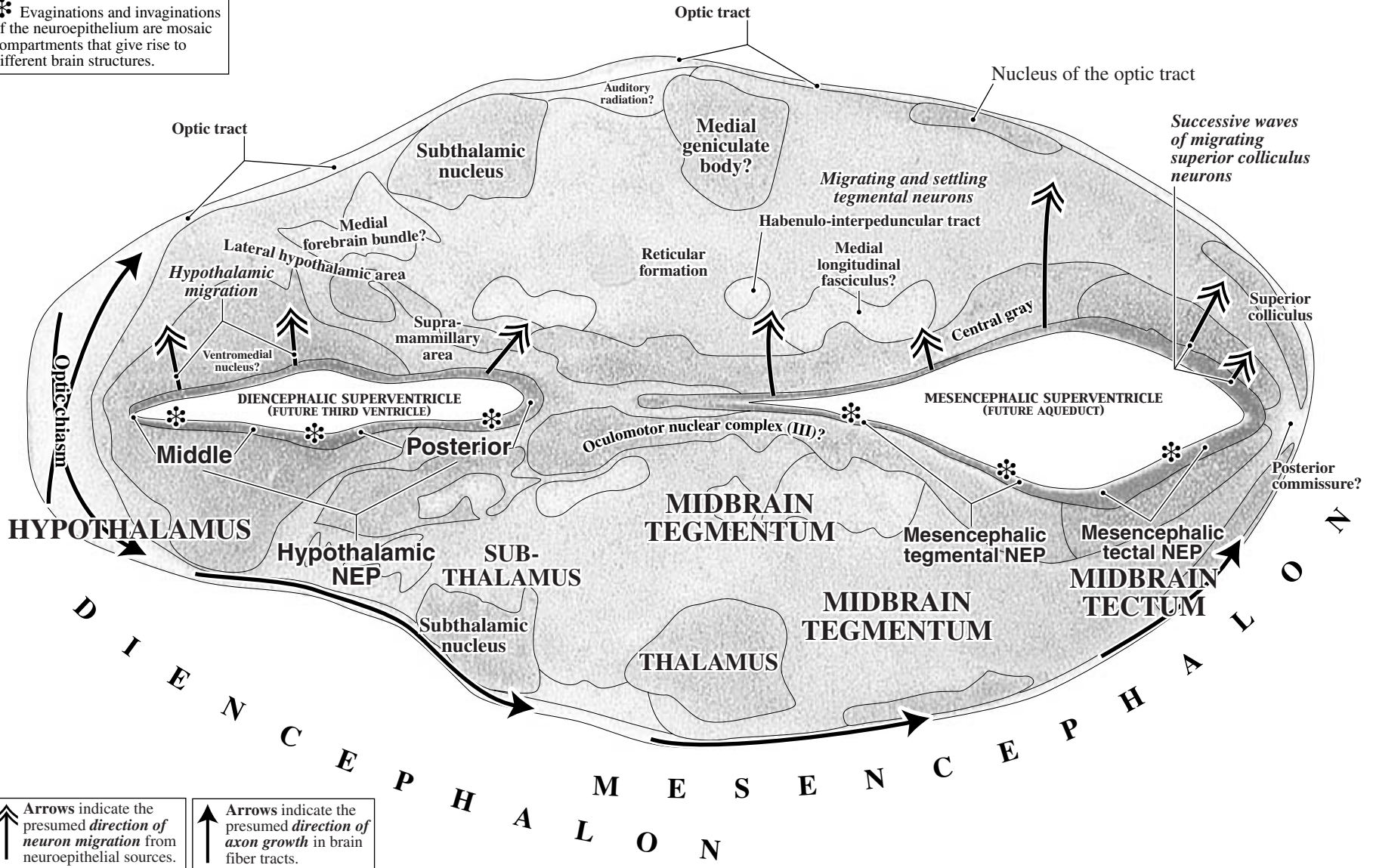
See the entire section in parts A and B of this plate on the preceding pages.

PLATE 92D

NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 93A

**GW9 Horizontal
CR 43 mm, C886
Level 11: Section 90**

**See the brain core enlarged
in parts C and D of this plate
on the following pages.**

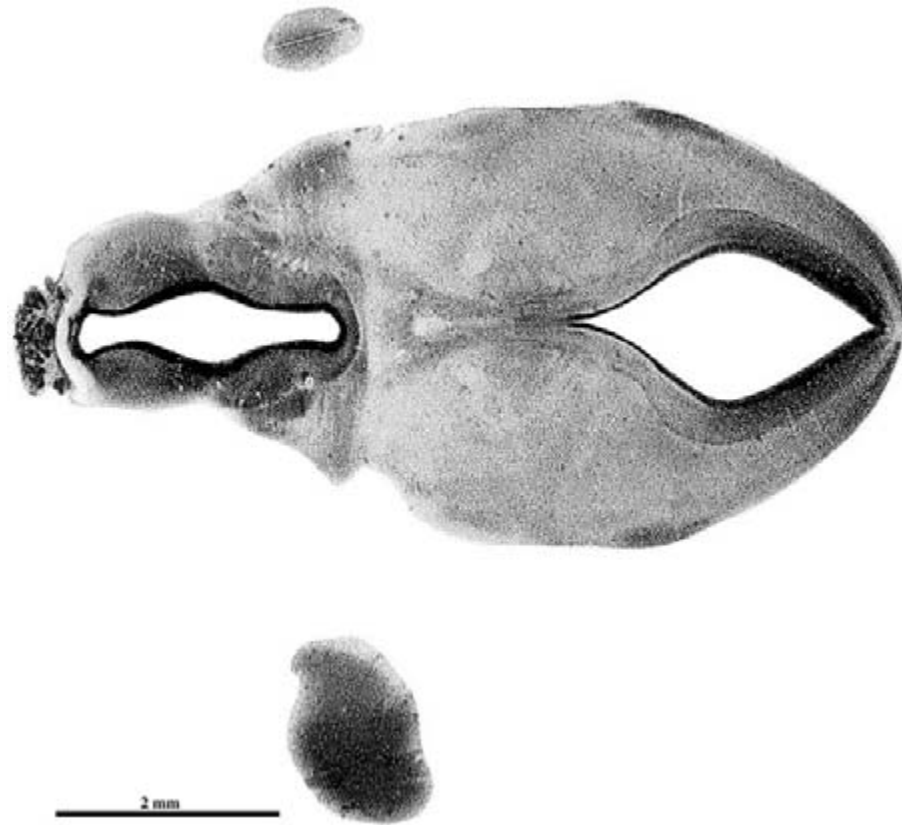
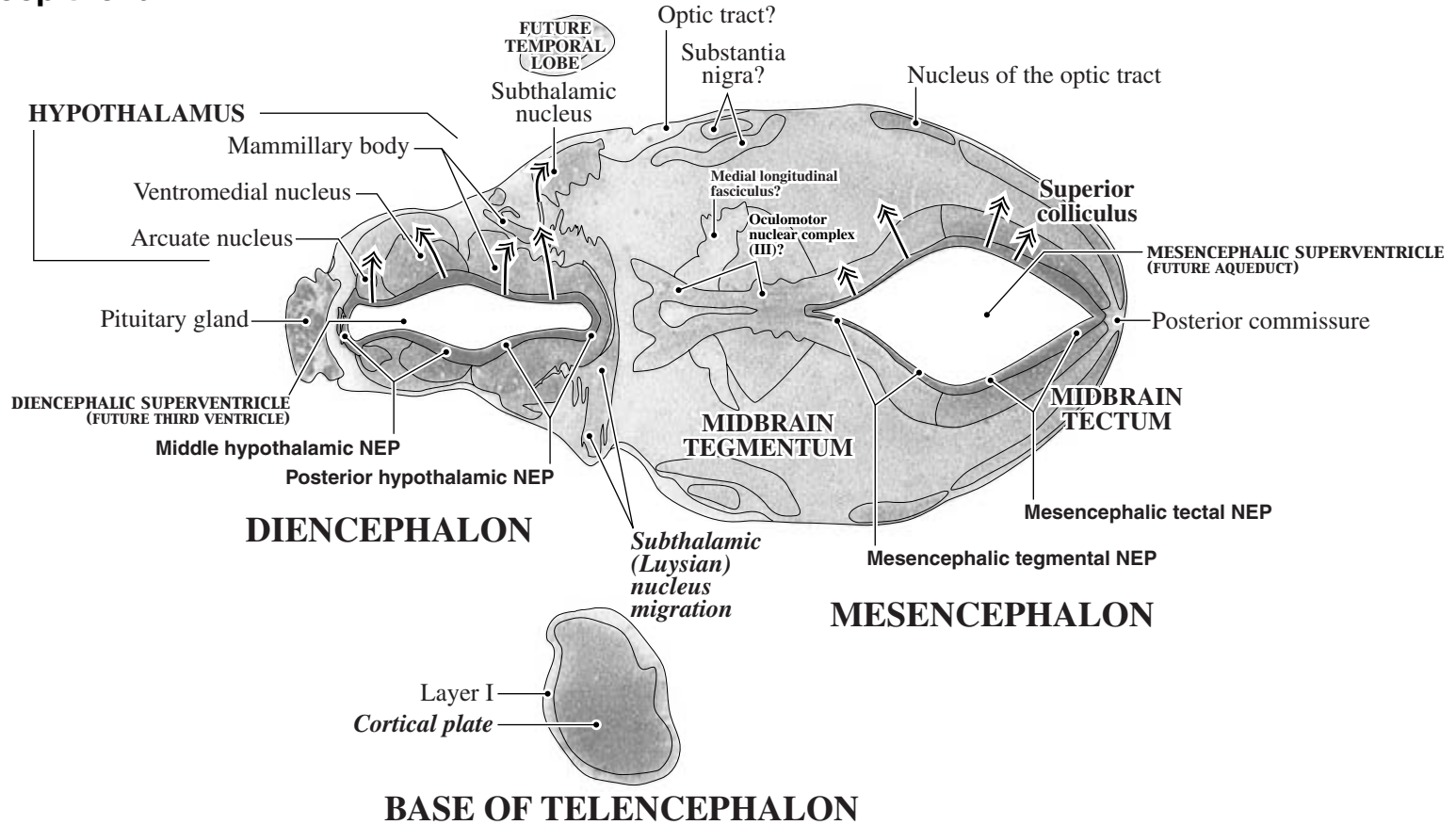


PLATE 93B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

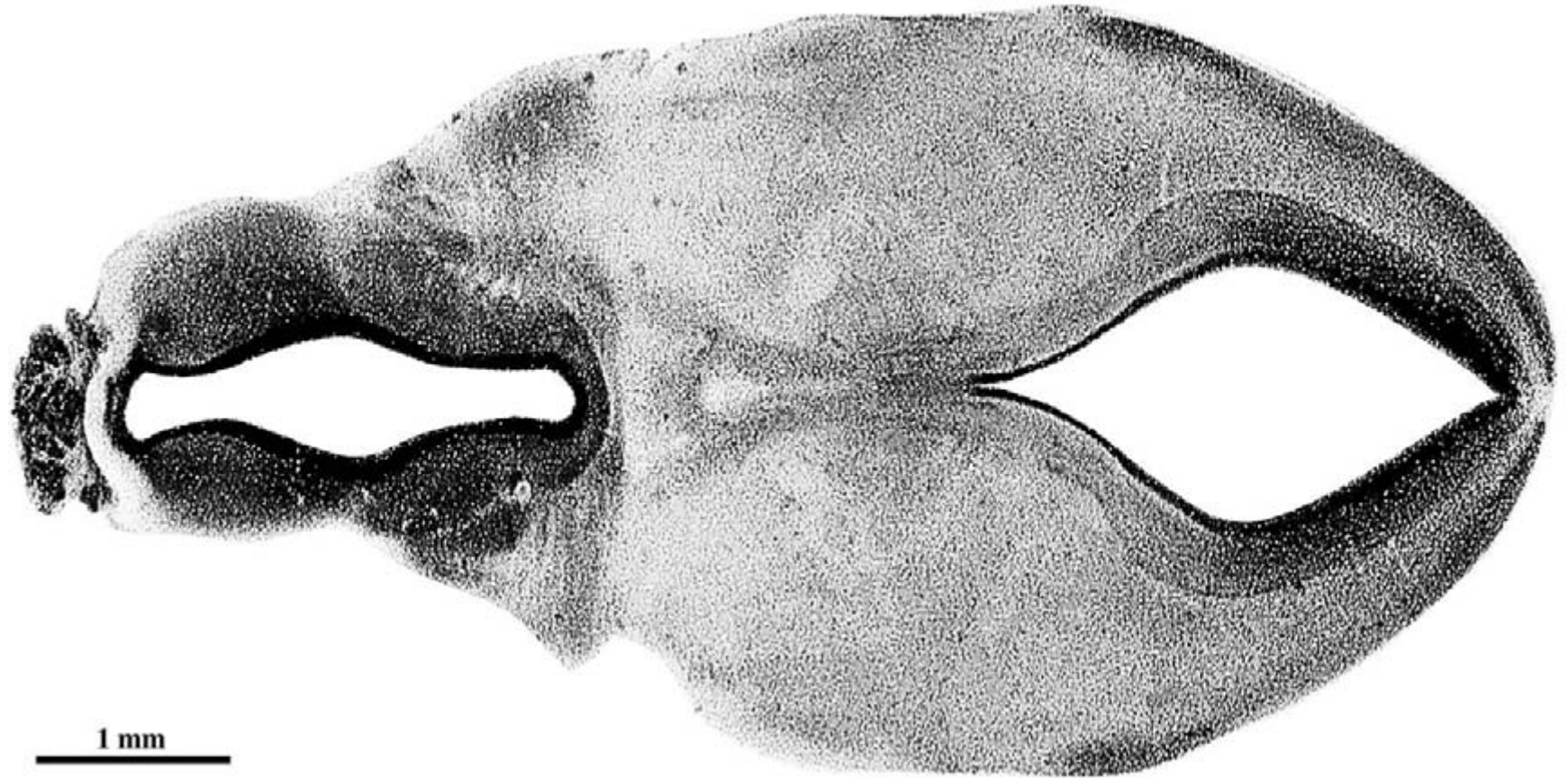
NEP - Neuroepithelium



↑ **Arrows** indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 93C

**GW9 Horizontal
CR 43 mm, C886
Level 11: Section 90**



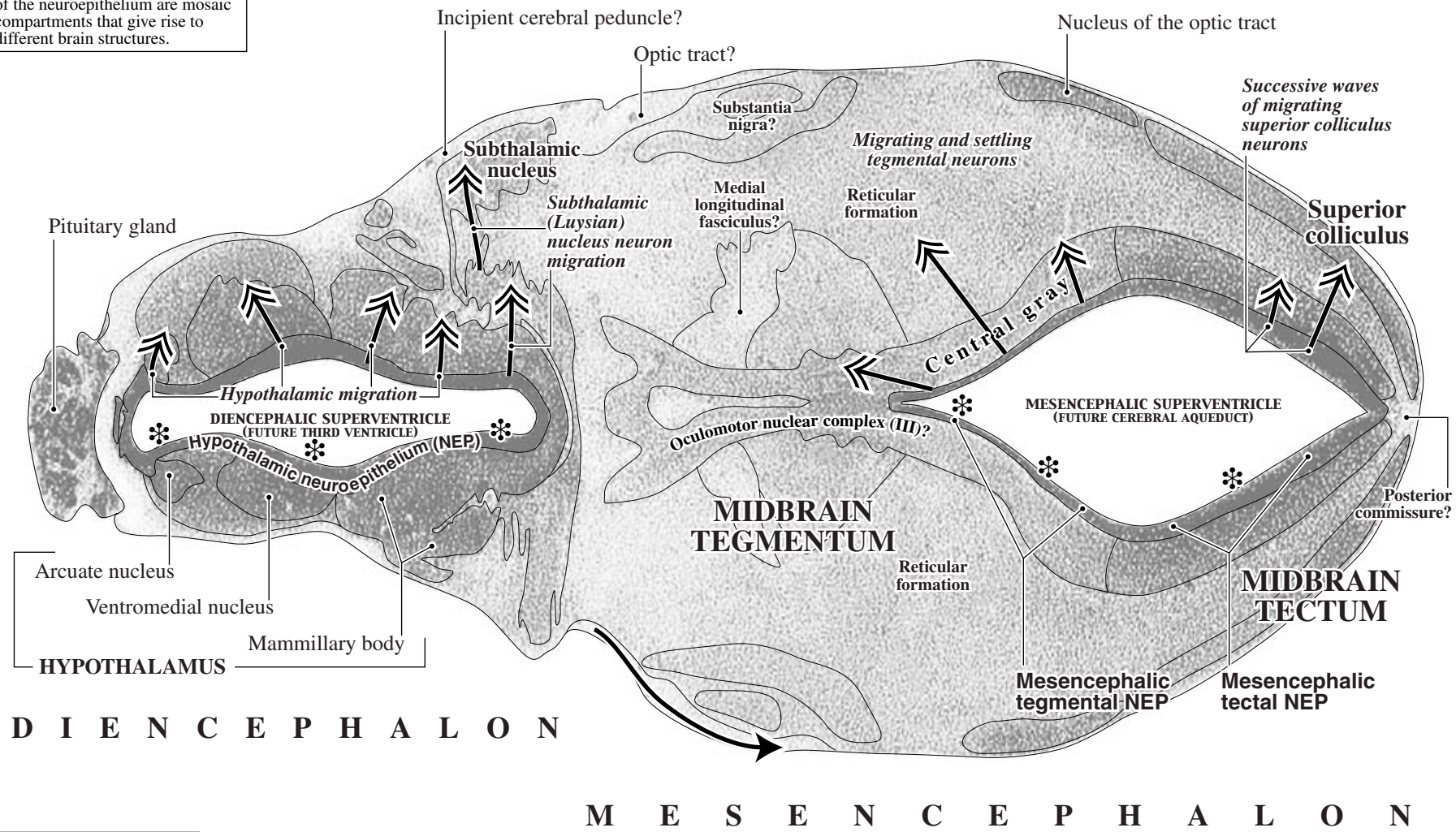
See the entire section in parts A and B of this plate on the preceding pages.

PLATE 93D

NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

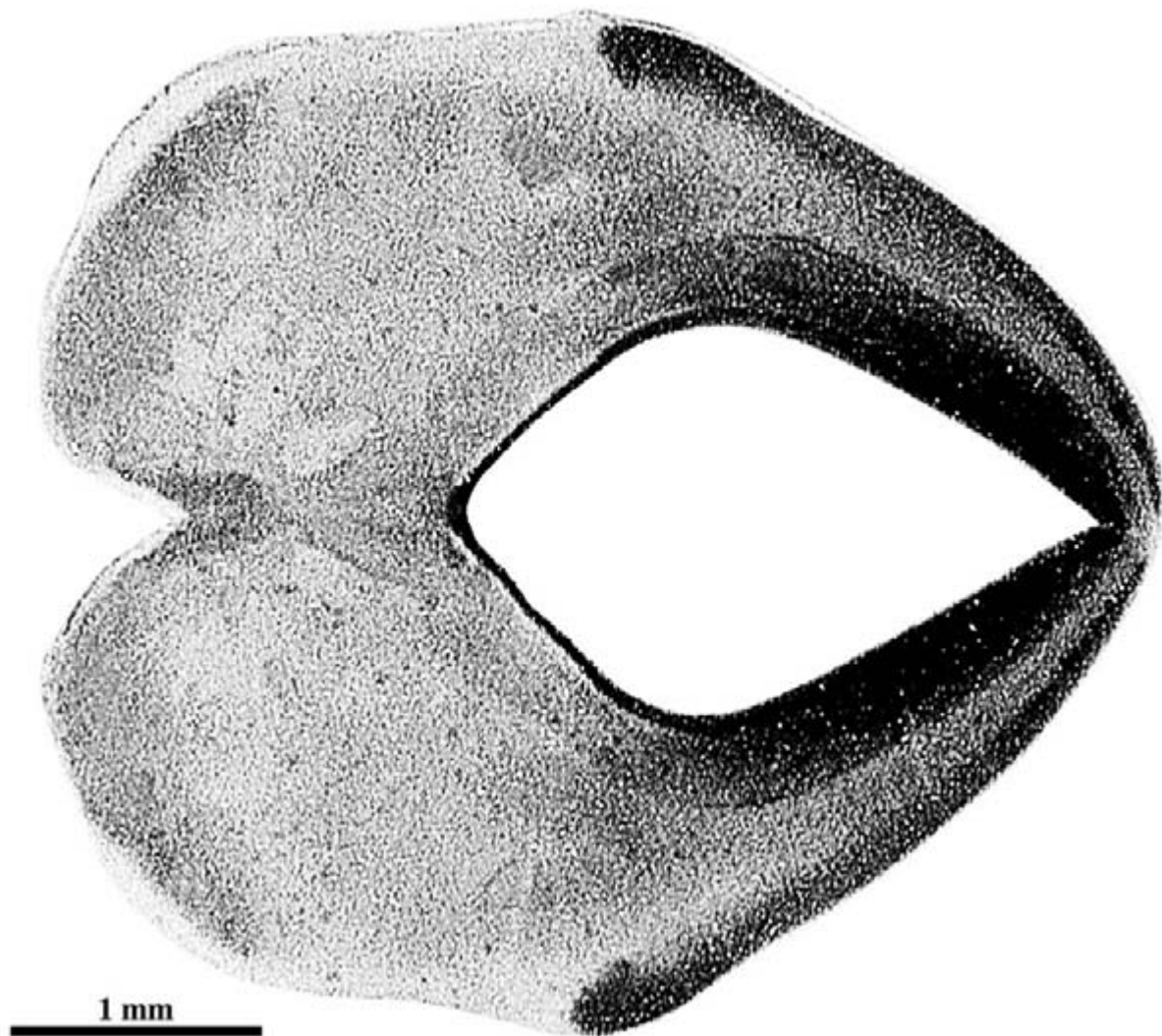


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 94A

**GW9 Horizontal
CR 43 mm, C886
Level 12: Section 100**



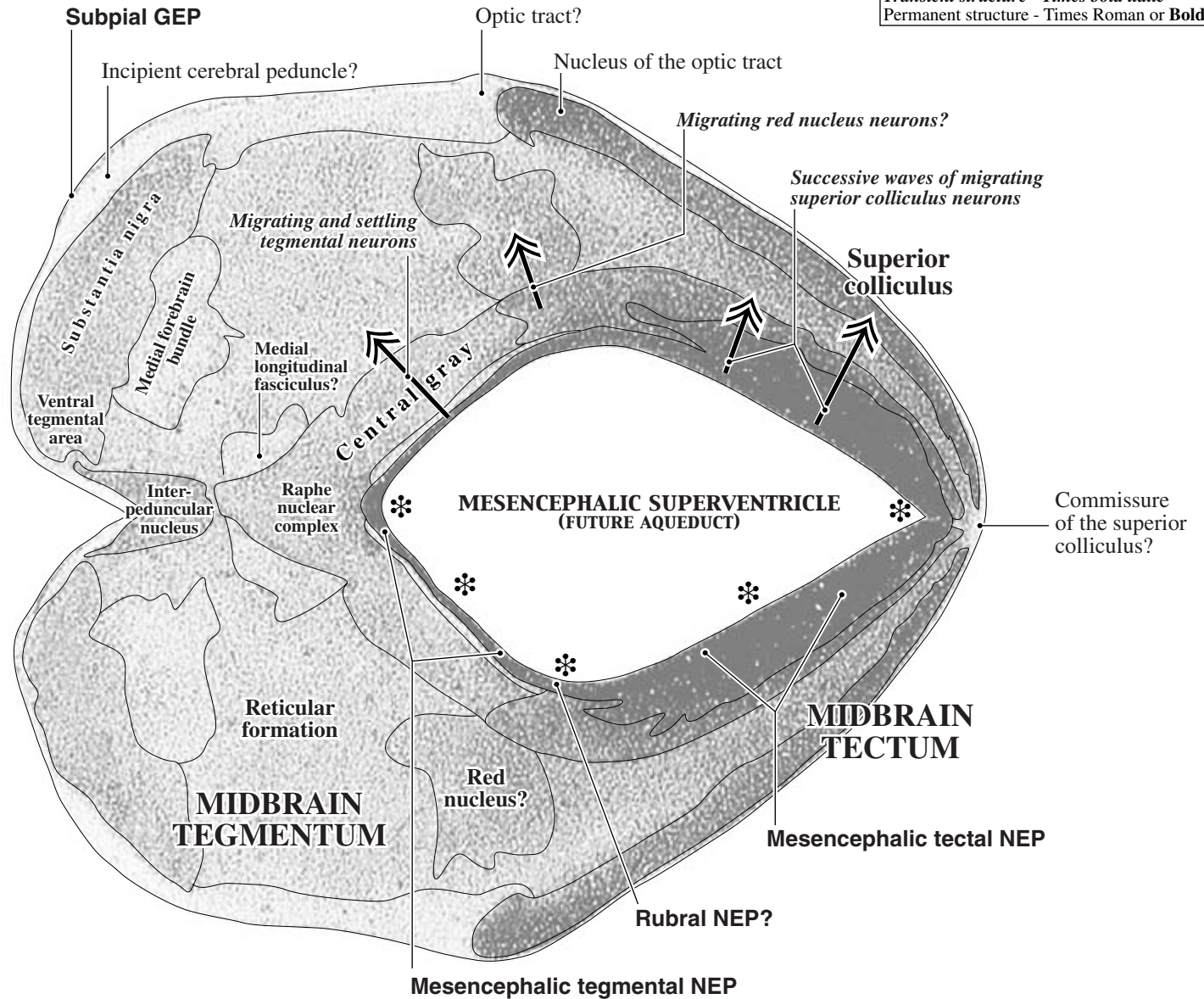
Levels 12 to 19 are shown only at higher magnification.

PLATE 94B

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

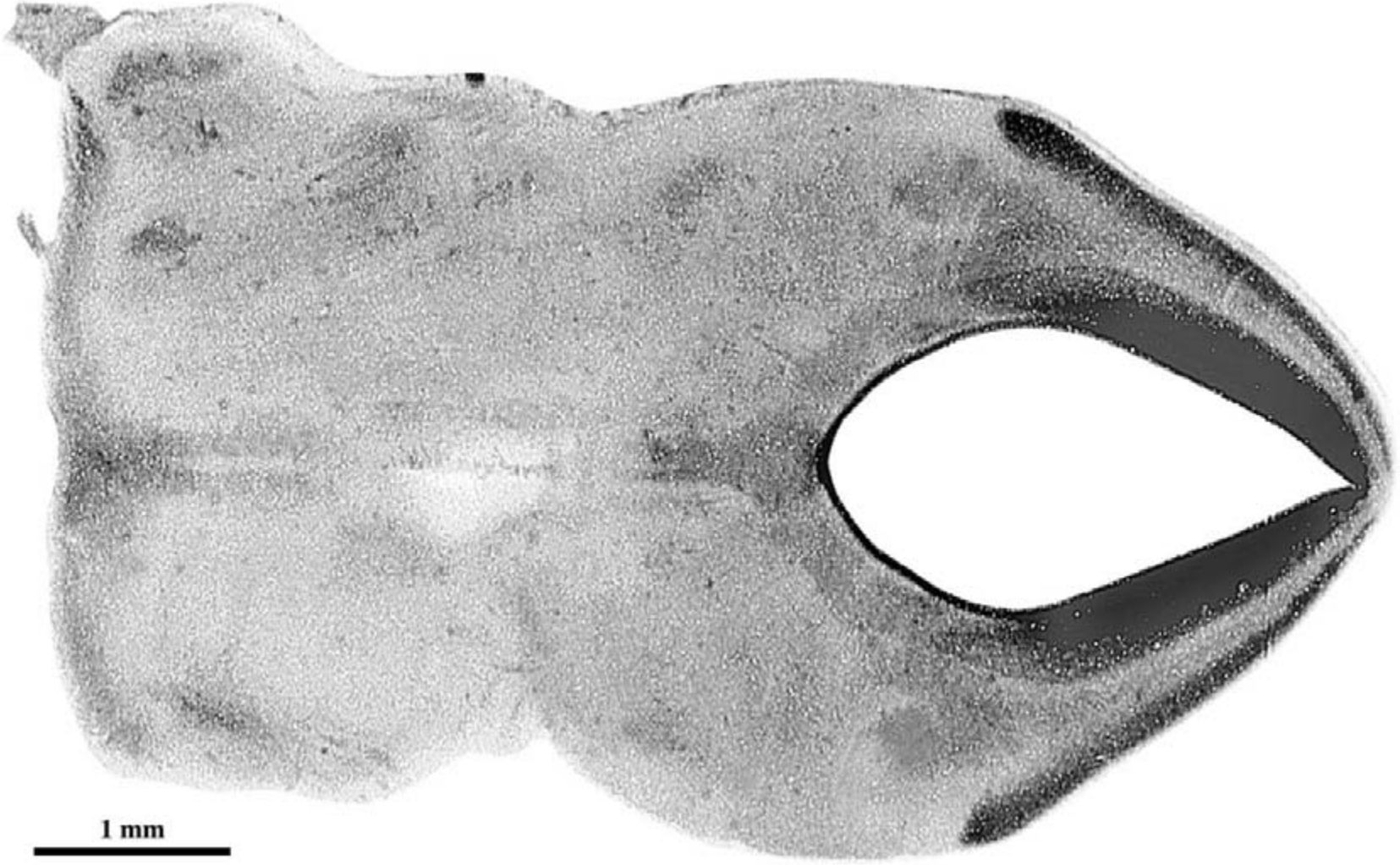
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 95A

**GW9 Horizontal
CR 43 mm, C886
Level 13: Section 105**



Levels 12 to 19 are shown only at higher magnification.

PLATE 95B

<p>* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.</p>	<p>ABBREVIATIONS: GEP - Glioepithelium NEP - Neuroepithelium</p>	<p>FONT KEY: VENTRICULAR DIVISIONS - CAPITALS Germinal zone - Helvetica bold <i>Transient structure - Times bold italic</i> Permanent structure - Times Roman or Bold</p>
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↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

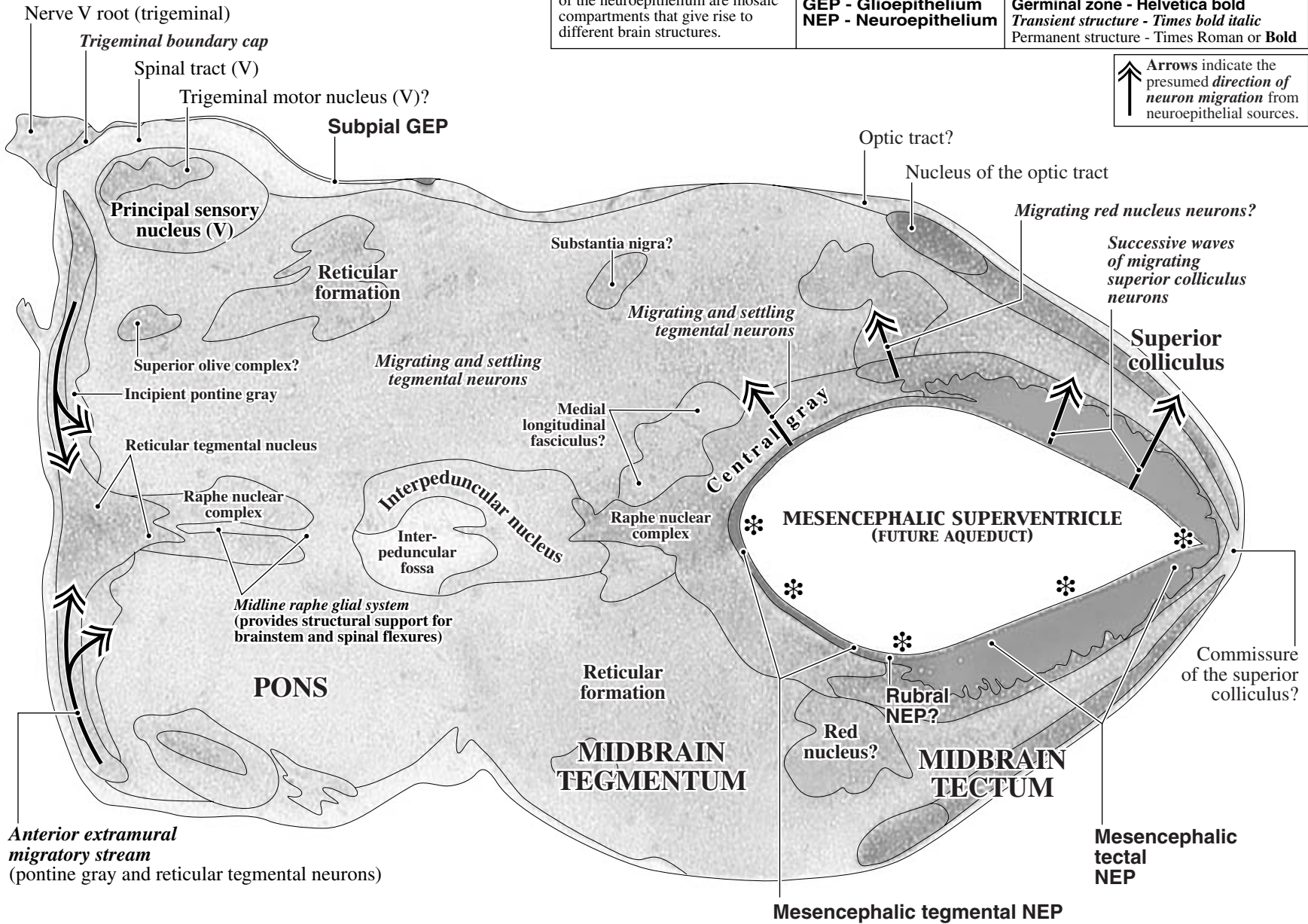
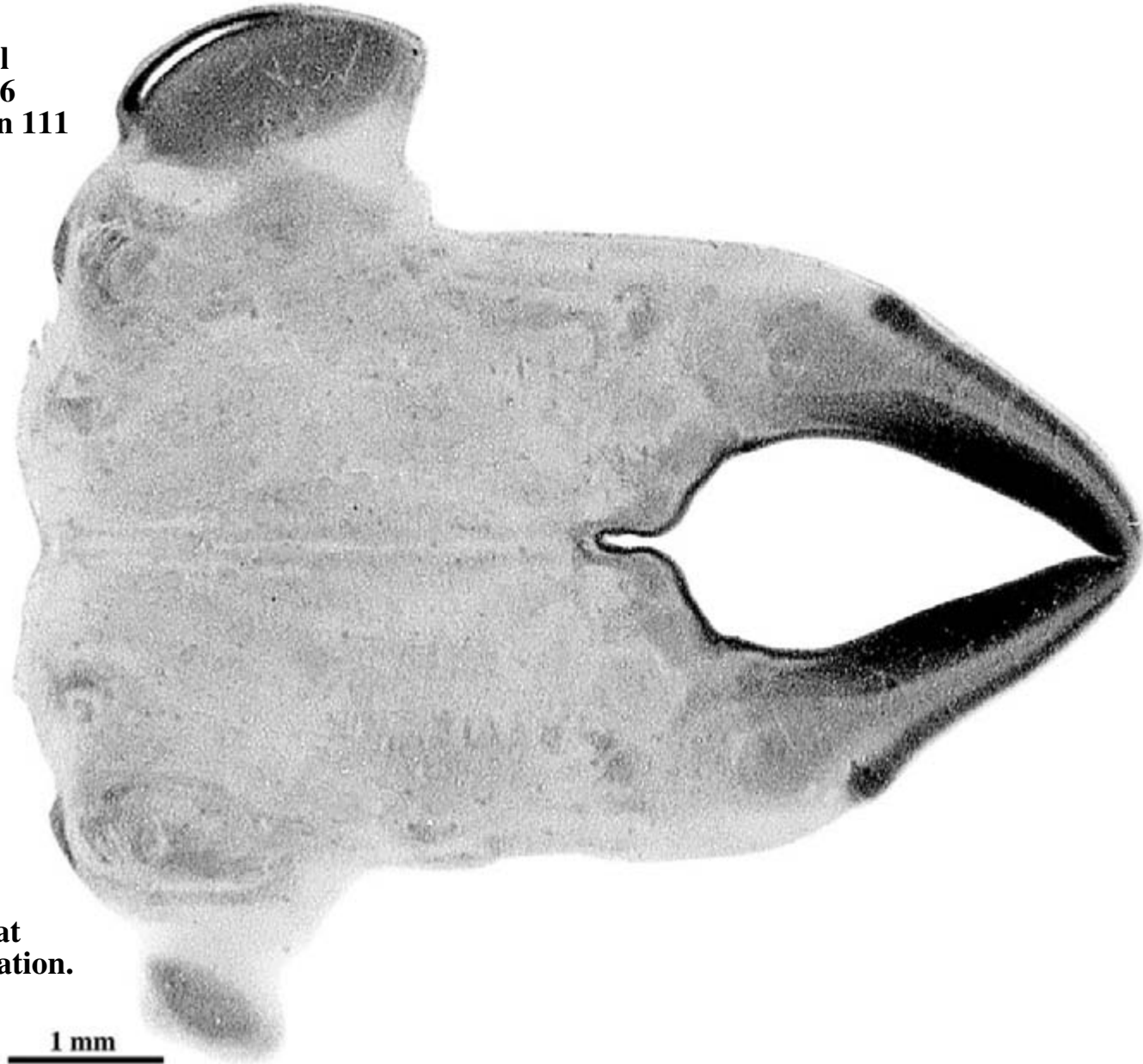


PLATE 96A

**GW9 Horizontal
CR 43 mm, C886
Level 14: Section 111**



**Levels 12 to 19
are shown only at
higher magnification.**

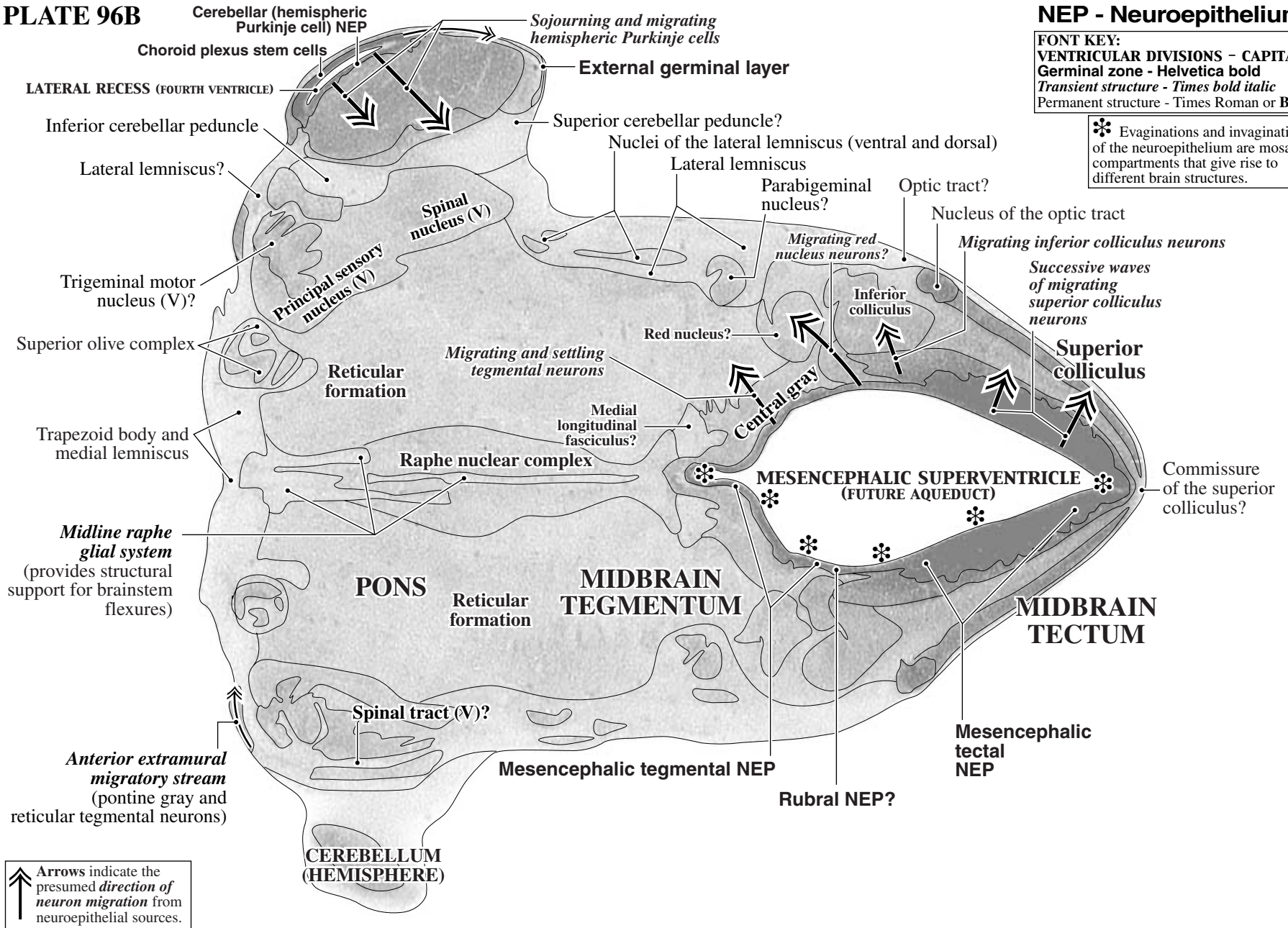
1 mm

PLATE 96B

NEP - Neuroepithelium

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

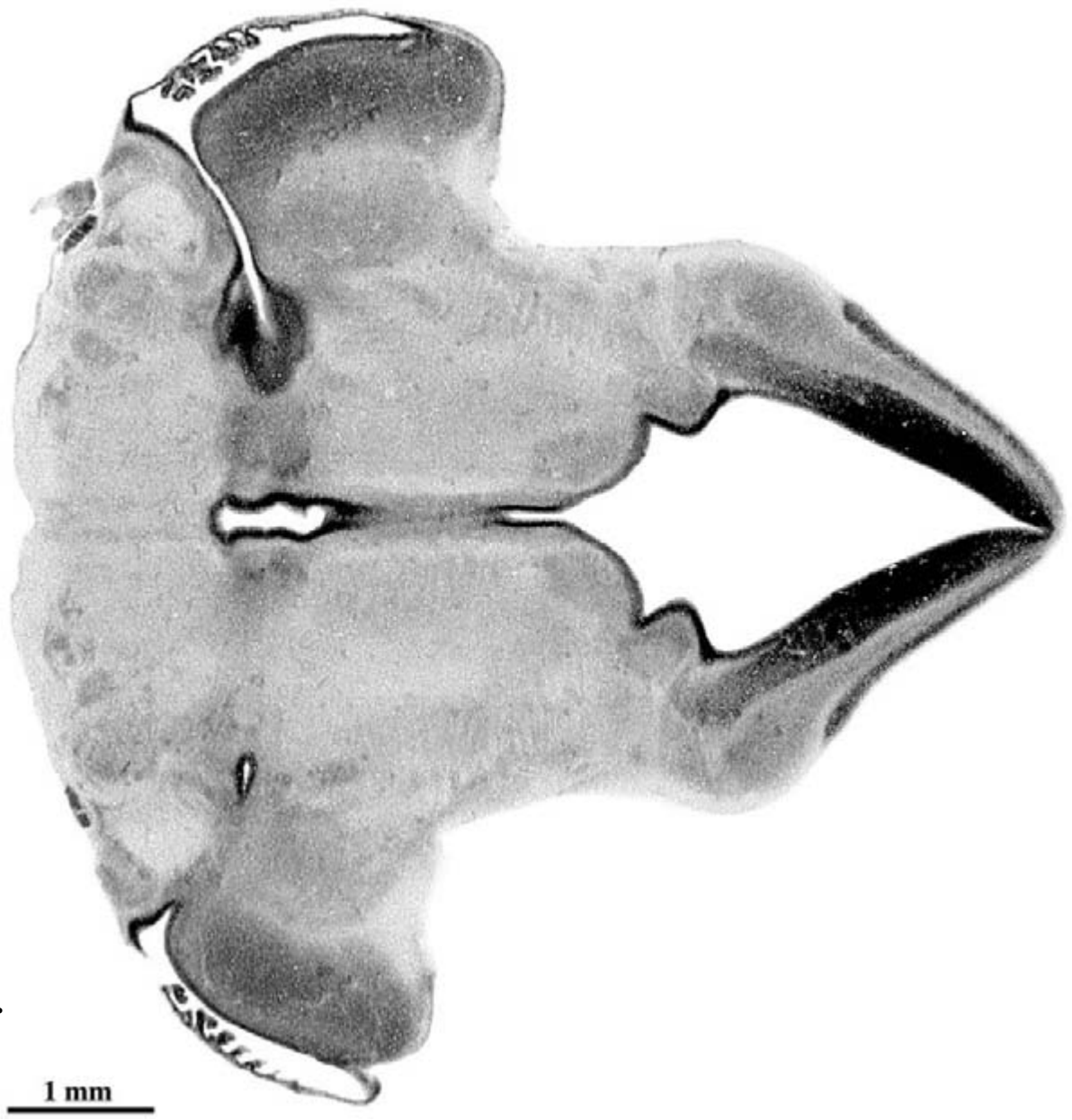
***** Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 97A

**GW9 Horizontal
CR 43 mm, C886
Level 15: Section 117**



**Levels 12 to 19
are shown only at
higher magnification.**

1 mm

PLATE 97B

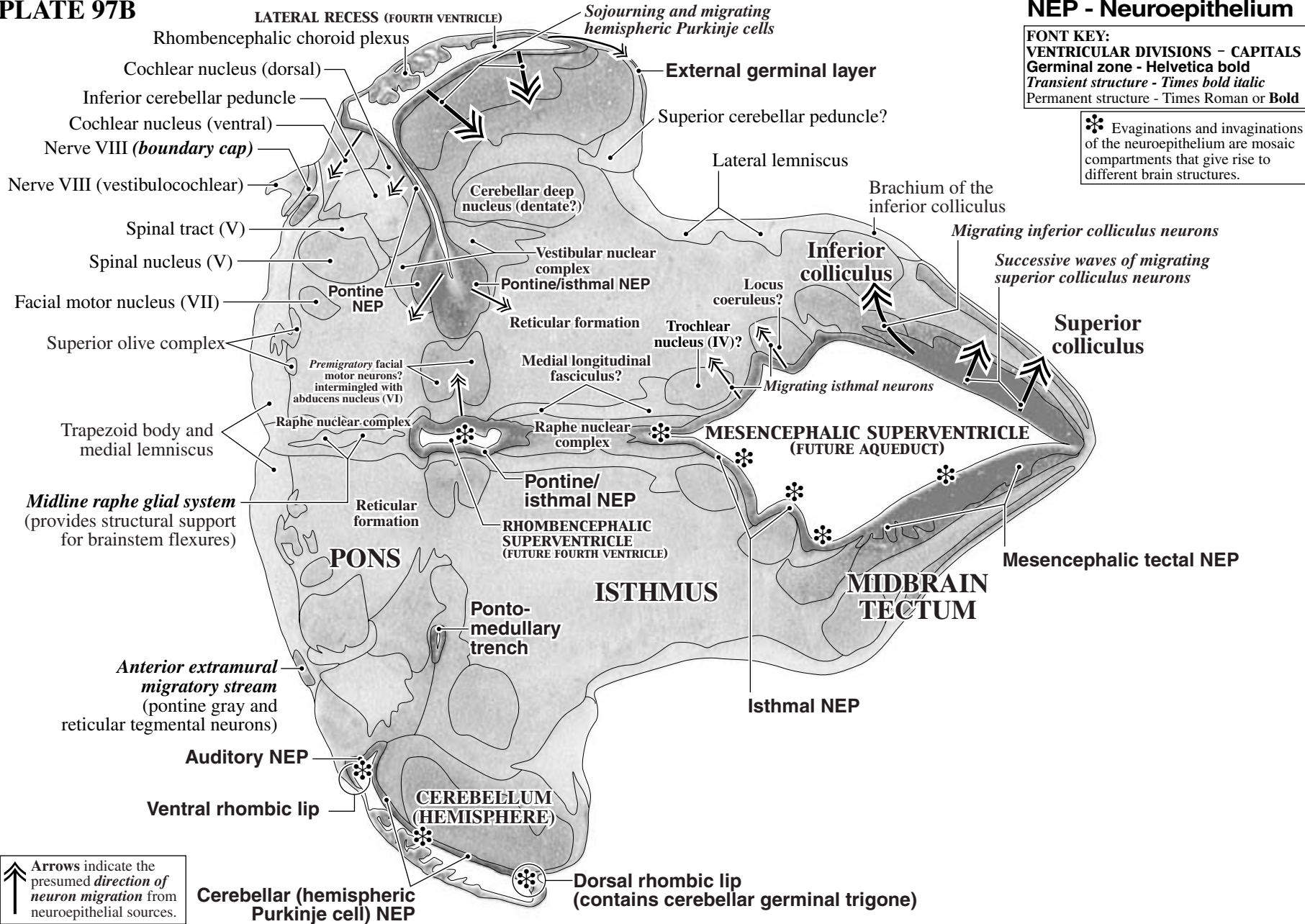
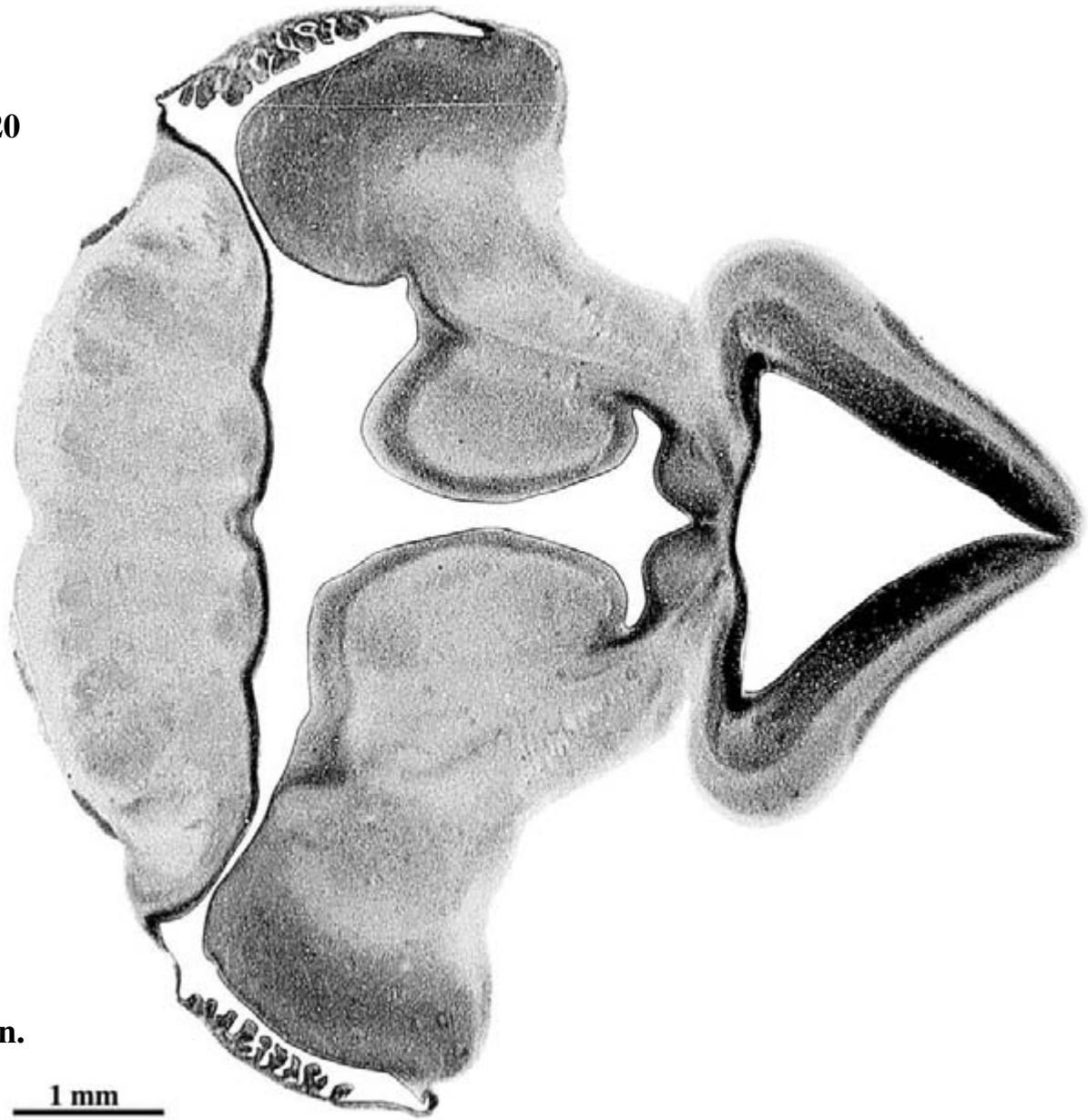


PLATE 98A

**GW9 Horizontal
CR 43 mm, C886
Level 16: Section 120**



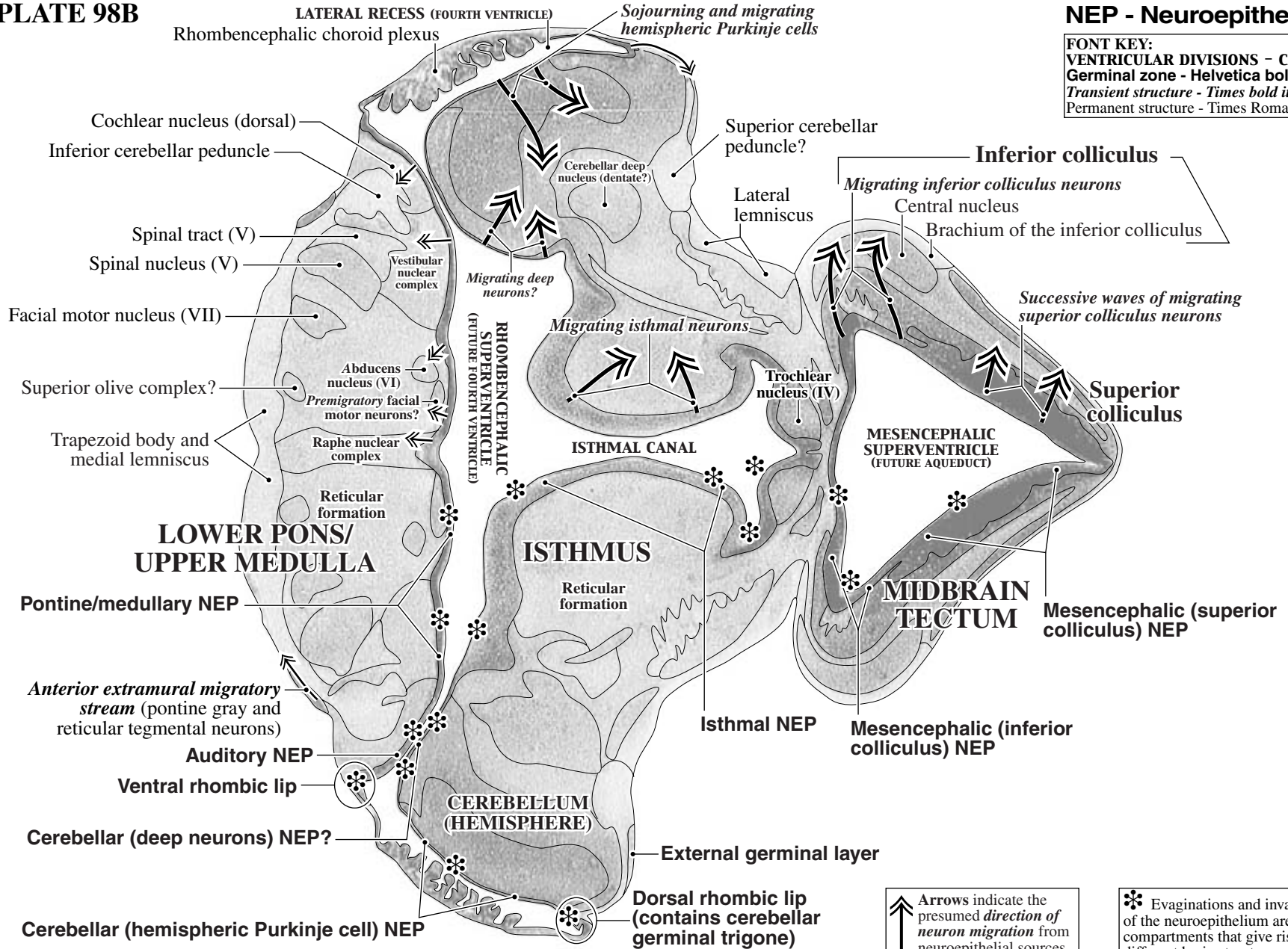
**Levels 12 to 19
are shown only at
higher magnification.**

1 mm

PLATE 98B

NEP - Neuroepithelium

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

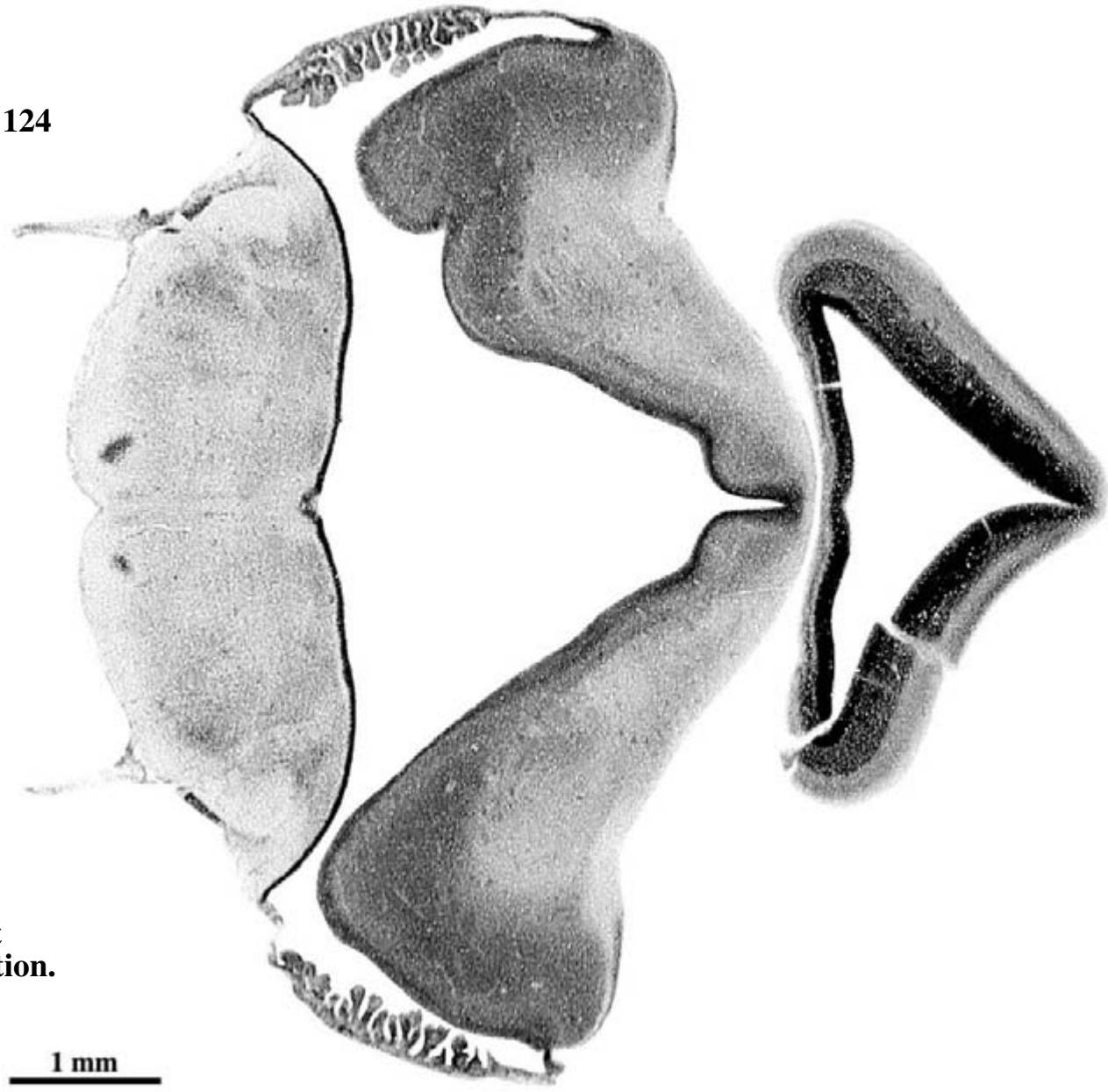


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 99A

**GW9 Horizontal
CR 43 mm, C886
Level 17: Section 124**



**Levels 12 to 19
are shown only at
higher magnification.**

1 mm

PLATE 99B

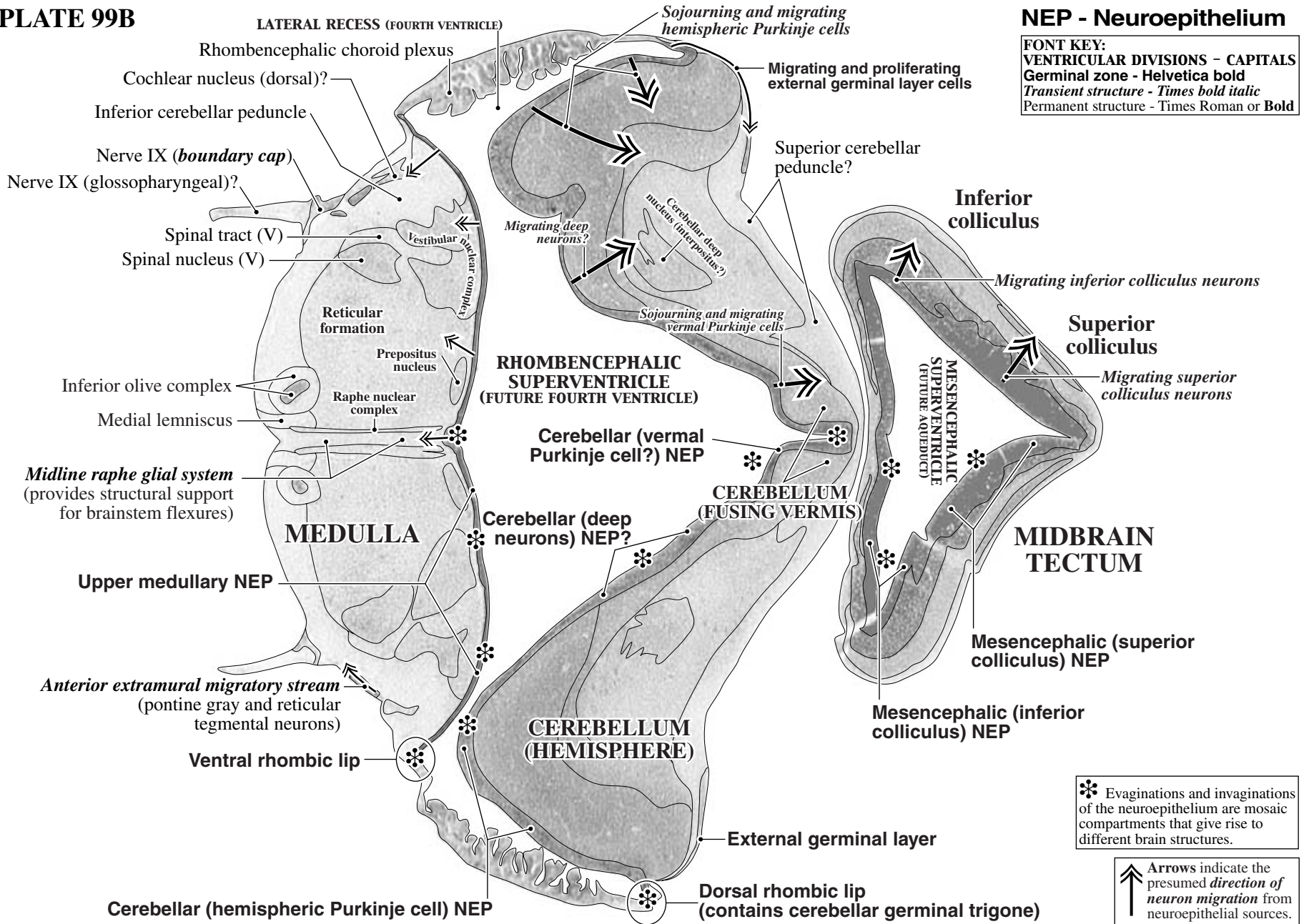
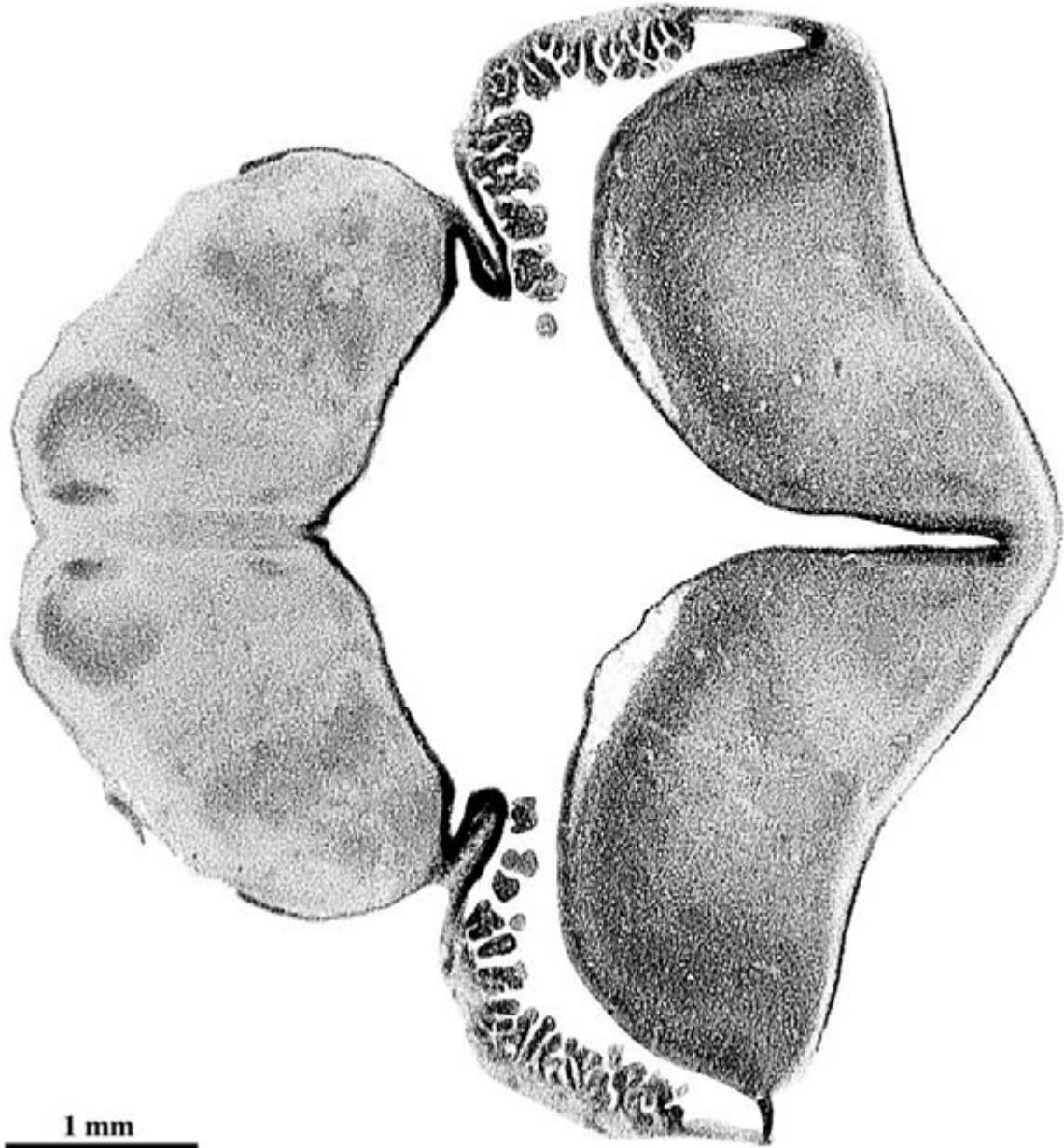


PLATE 100A

**GW9 Horizontal
CR 43 mm, C886
Level 18: Section 134**



**Levels 12 to 19
are shown only at
higher magnification.**

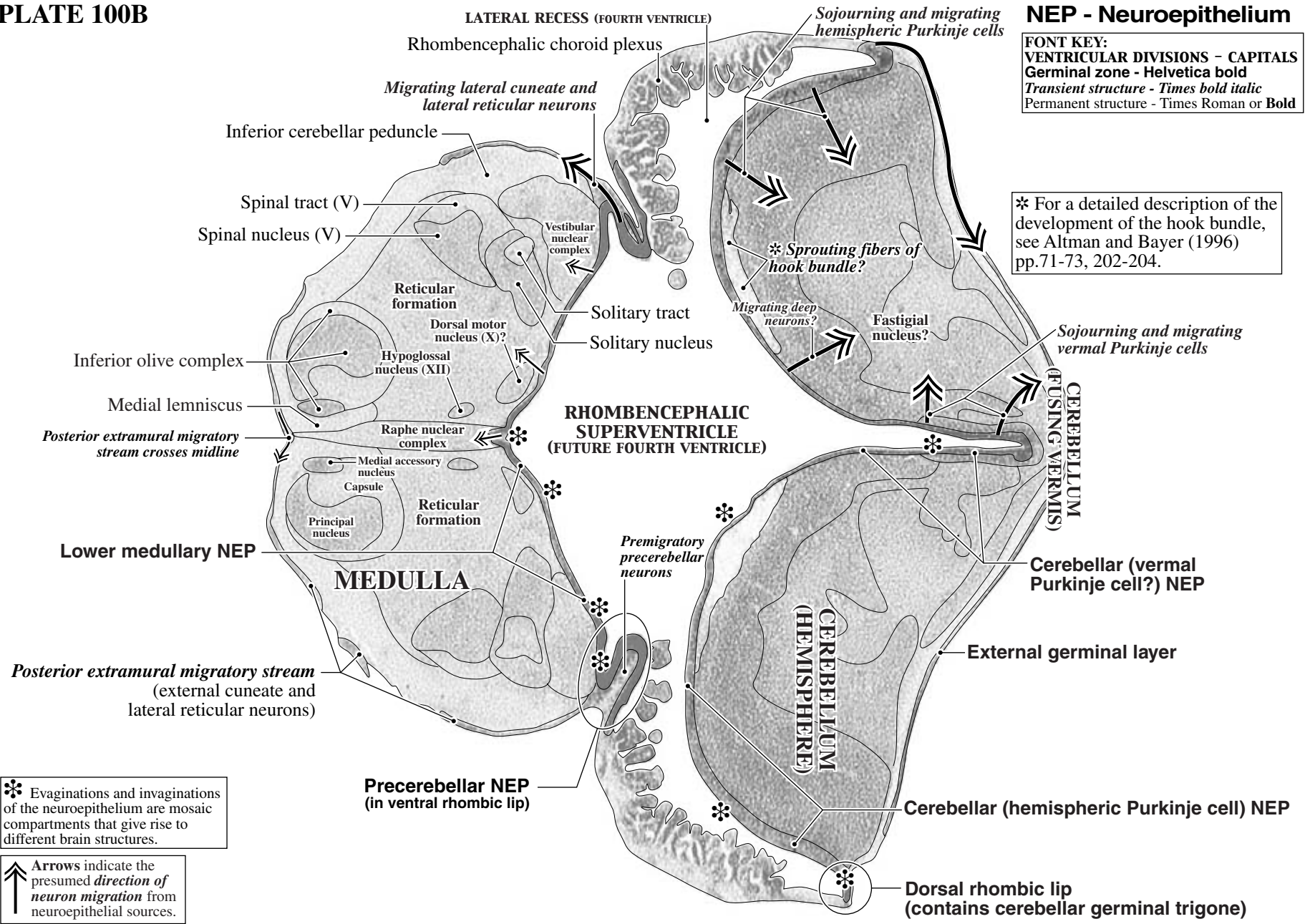
1 mm

PLATE 100B

NEP - Neuroepithelium

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

** For a detailed description of the development of the hook bundle, see Altman and Bayer (1996) pp.71-73, 202-204.*

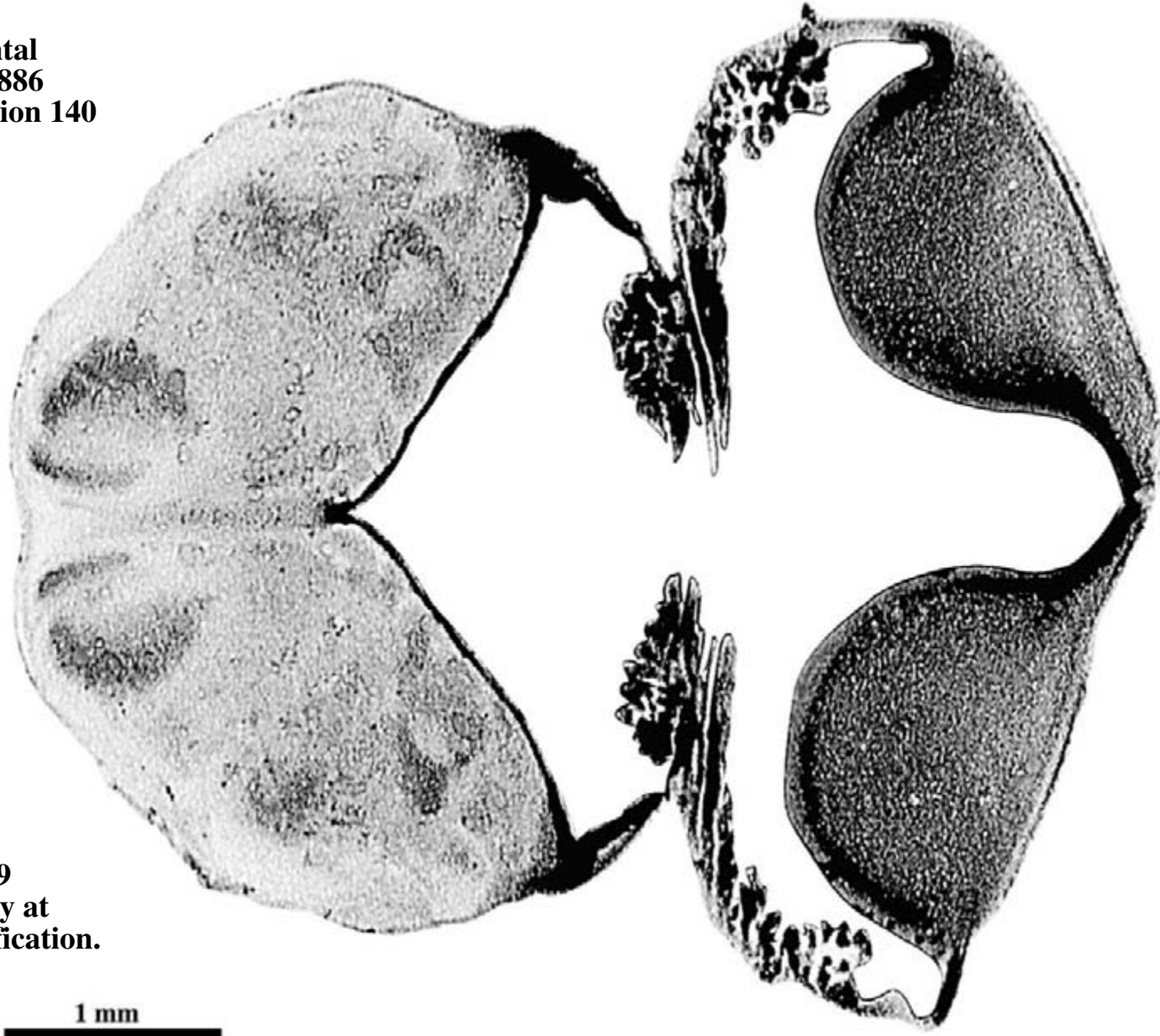


* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 101A

**GW9 Horizontal
CR 43 mm, C886
Level 19: Section 140**



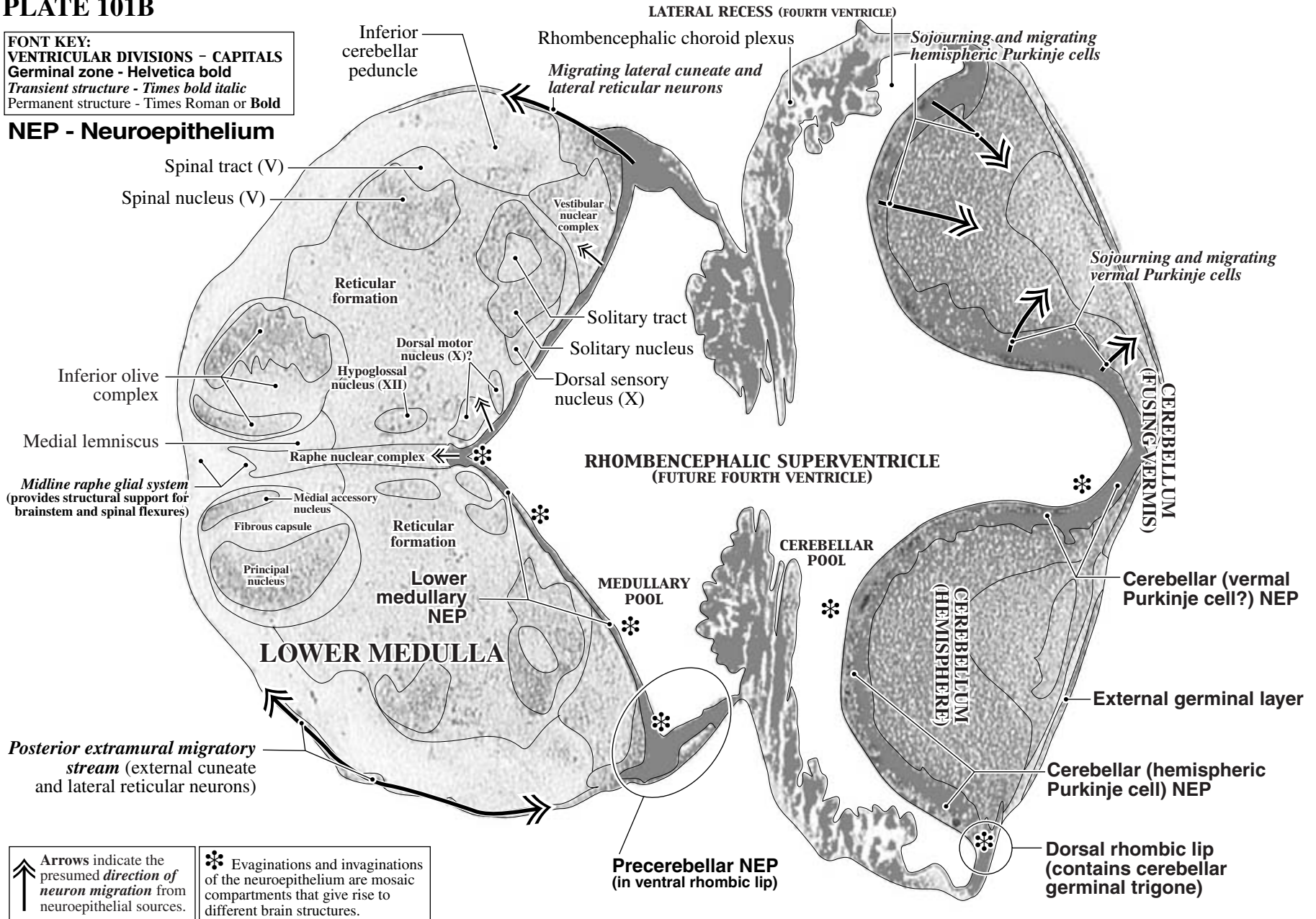
**Levels 12 to 19
are shown only at
higher magnification.**

1 mm

PLATE 101B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

NEP - Neuroepithelium



PART VI: GW9 CORONAL

This specimen is human fetus number 282 with a crown-rump length (CR) of 42 mm estimated to be at gestational week (GW) 9 (Minot Collection histological record number 841, referred to here as M841). The fetus was embedded in paraffin, cut in 10- μ m thick sections, and stained with borax carmine and Lyon's blue. No information is available on date of collection (sometime between 1900 and 1910) and the kind of fixative. Since there is no photograph of this brain before it was embedded and cut, a specimen from Hochstetter (1919) that is comparable to M841 has been modified to show the approximate section plane and external features of the brain at GW9 (inset, **Figure 5**). Like most of the specimens in this Volume, the sections are not cut exactly in one plane; M841 is midway between coronal and horizontal. Since the cerebral cortex is in every section and the brainstem is cut in a more horizontal orientation, the brain more closely resembles a coronally sectioned brain. Photographs of 22 sections (**Levels 1-22**) are illustrated at low magnification in **Plates 102-121**. Excellent tissue detail is preserved at low magnification, but there is a fine granular precipitate visible at high magnification.

M841 is similar to the other GW9 specimens in the level of brain maturation. The chief reason for including this specimen is to provide a third perspective for viewing brain structure at GW9. In the cerebral cortex, the *neuroepithelium* is prominent as the sole germinal matrix; the *stratified transitional field (STF)* consists of *STF1*, *STF5*, and *STF4* only in lateral areas. The anterolateral (thicker) to dorsomedial (thinner) maturation gradient in the cortical plate and *STF* layers are evident. In this specimen, the olfactory evagination is most evident coming from the basal telencephalic rather than from the cerebral cortical neuroepithelium. In anterolateral parts of the cerebral cortex, streams of neurons and glia appear to leave *STF4* and enter the *lateral migratory stream*. A massive *neuroepithelium/subventricular zone* overlies the amyg-

dala, nucleus accumbens, and striatum (caudate and putamen) where neurons (and glia) are being generated.

The cerebellum has a definite *neuroepithelium* at the ventricular surface. Most of the Purkinje cells are sojourning in a thick dense layer outside the neuroepithelium, and others are migrating upward. Many of the deep neurons are superficial in the cerebellum, but some are migrating downward to intermingle with upwardly migrating Purkinje cells. The cortical surface is partially covered by an *external germinal layer (egl)* that is actively producing neuronal stem cells, as it grows over the surface of the cerebellar cortex.

The third ventricle, aqueduct, and fourth ventricle are lined by thin *neuroepithelia*. The midbrain tegmentum, pons, and medulla have the thinnest neuroepithelia indicating that only the latest generated neurons are being produced at this time. The thick precerebellar neuroepithelium is an exception in the medulla. Thicker neuroepithelia are in the cerebellum (see above) and midbrain tectum, indicating many neurons are still being generated, although the majority of the neurons in these sites are already postmitotic. The neuroepithelium is still thicker in the hypothalamus and thalamus, in accordance with the later maturation of the diencephalon compared to the rest of the brainstem.

Neurons throughout the diencephalon, midbrain tegmentum, pons, and medulla are migrating and settling. Nuclear divisions are very indistinct throughout the diencephalon because more neurons are migrating and have not yet settled permanently. More definition is seen in the midbrain tegmentum, pons, and medulla where cell migration is waning. As with the other GW9 specimens, the *anterior extramural* and *posterior extramural migratory streams* are dense subpial accumulations in the medulla and pons.

GW9 "CORONAL" SECTION PLANES

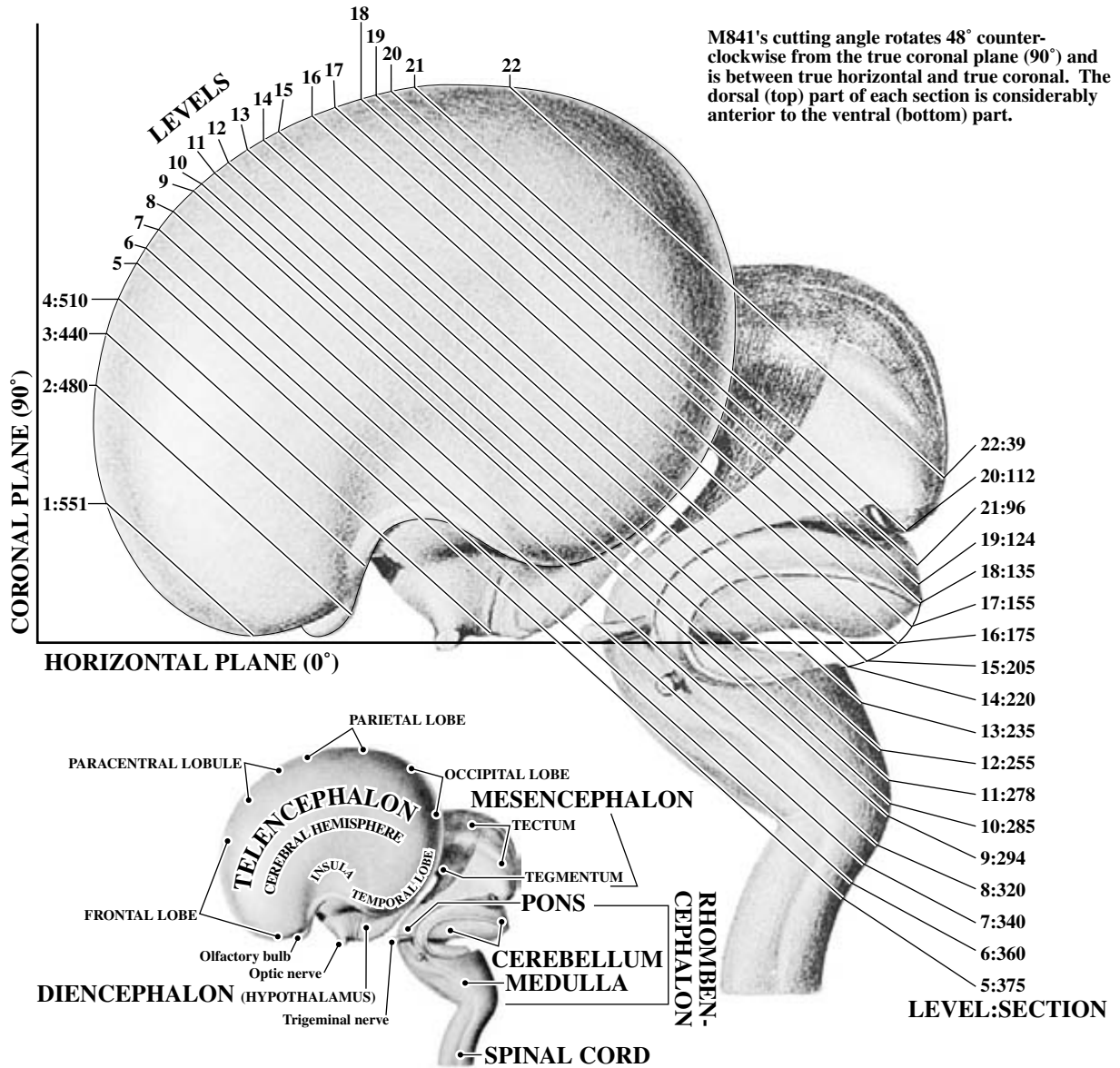
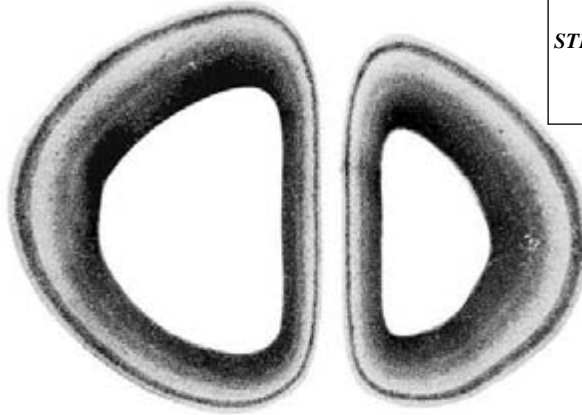


Figure 5. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 38 mm (modified from Figure 43, Table VII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of M841 in the following pages. The small inset identifies the major structural features. The line in the cerebellum and dorsal edges of the pons and medulla is the cut edge of the medullary velum.

PLATE 102A

GW9 Coronal
CR 42 mm
M841

Level 1: Section 551

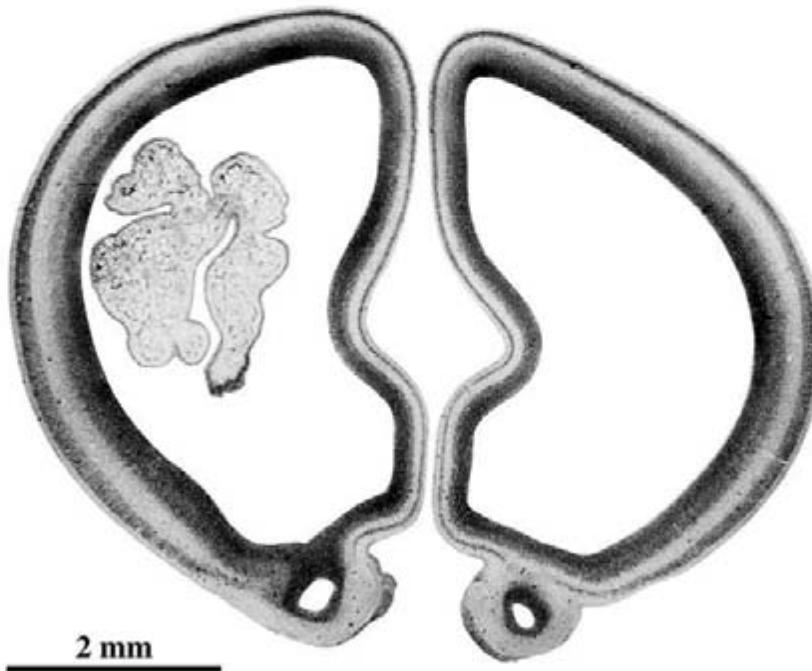


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

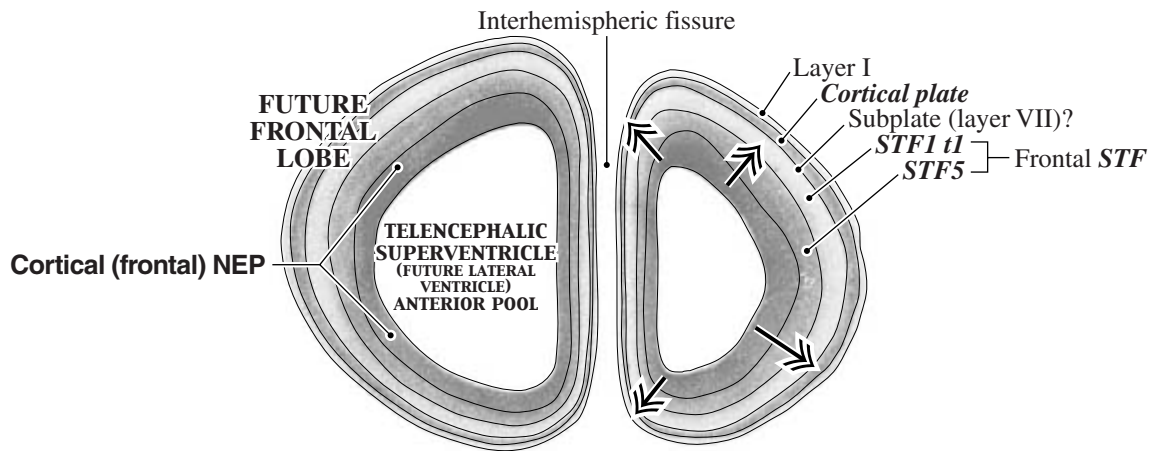
STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

Level 2: Section 480

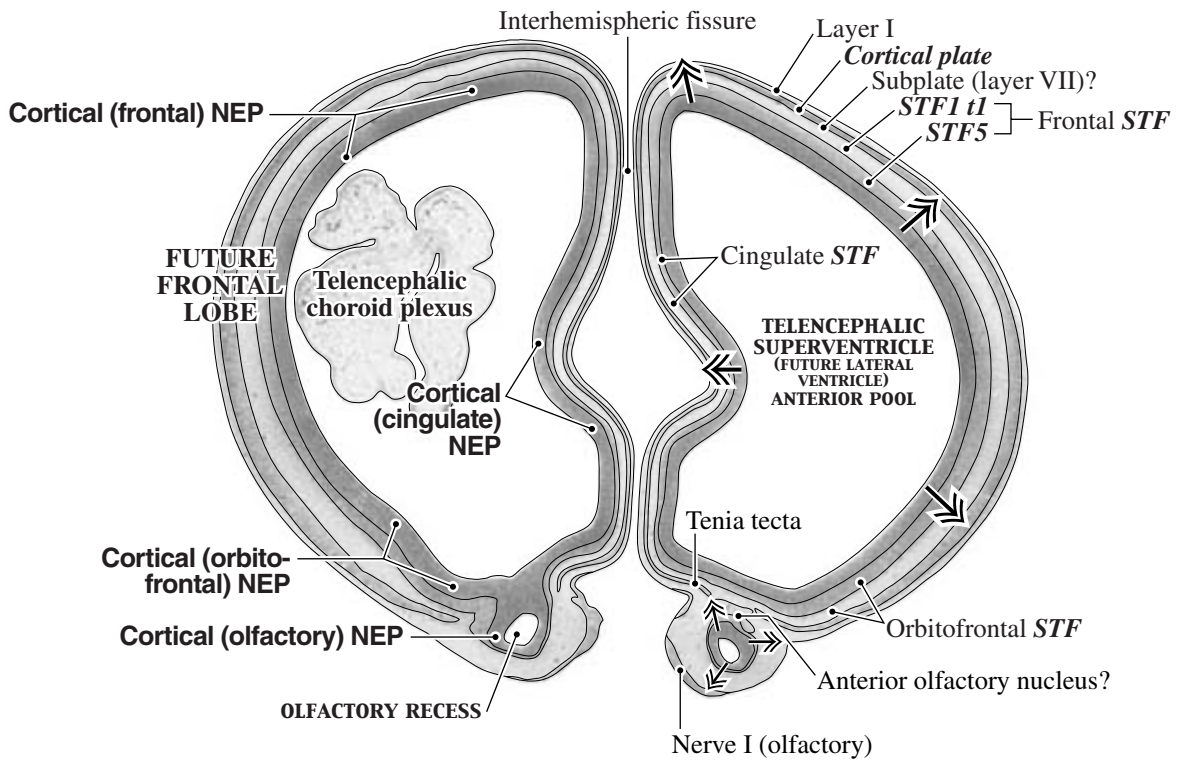


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

Level 1: Section 551



Level 2: Section 480



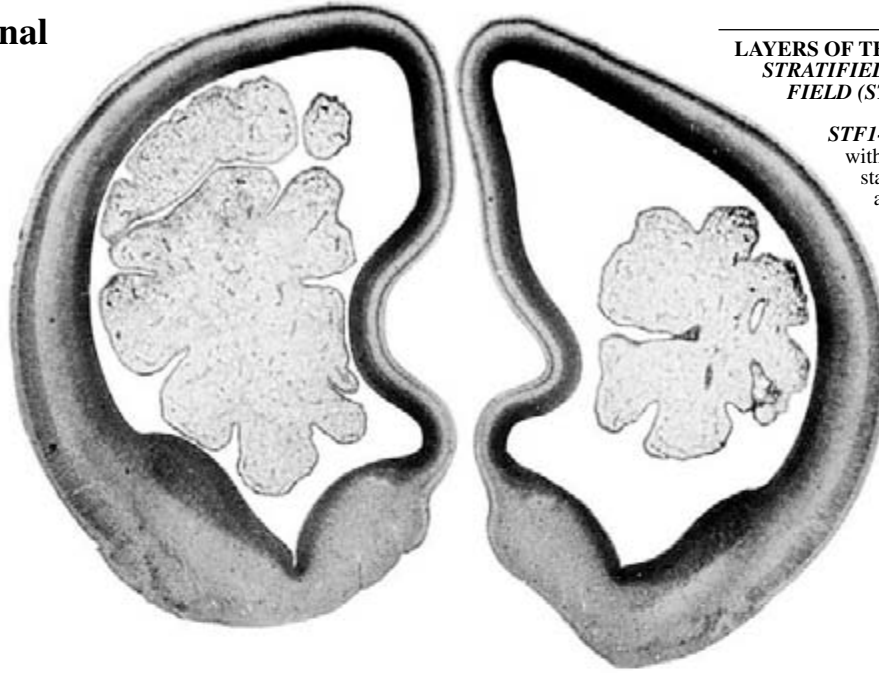
Cortical folding in the midline is a shrinkage artifact.

Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 103A

Level 3: Section 440

GW9 Coronal
CR 42 mm
M841

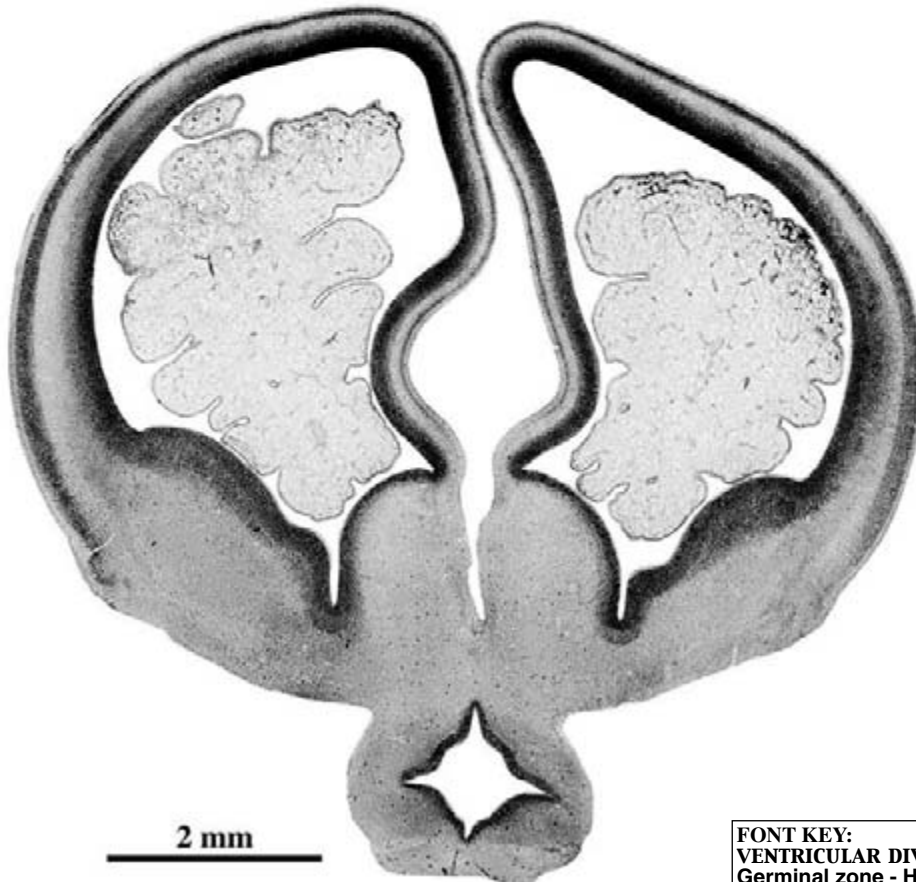


LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL
FIELD (STF)

STF1-Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal

Level 4: Section 410



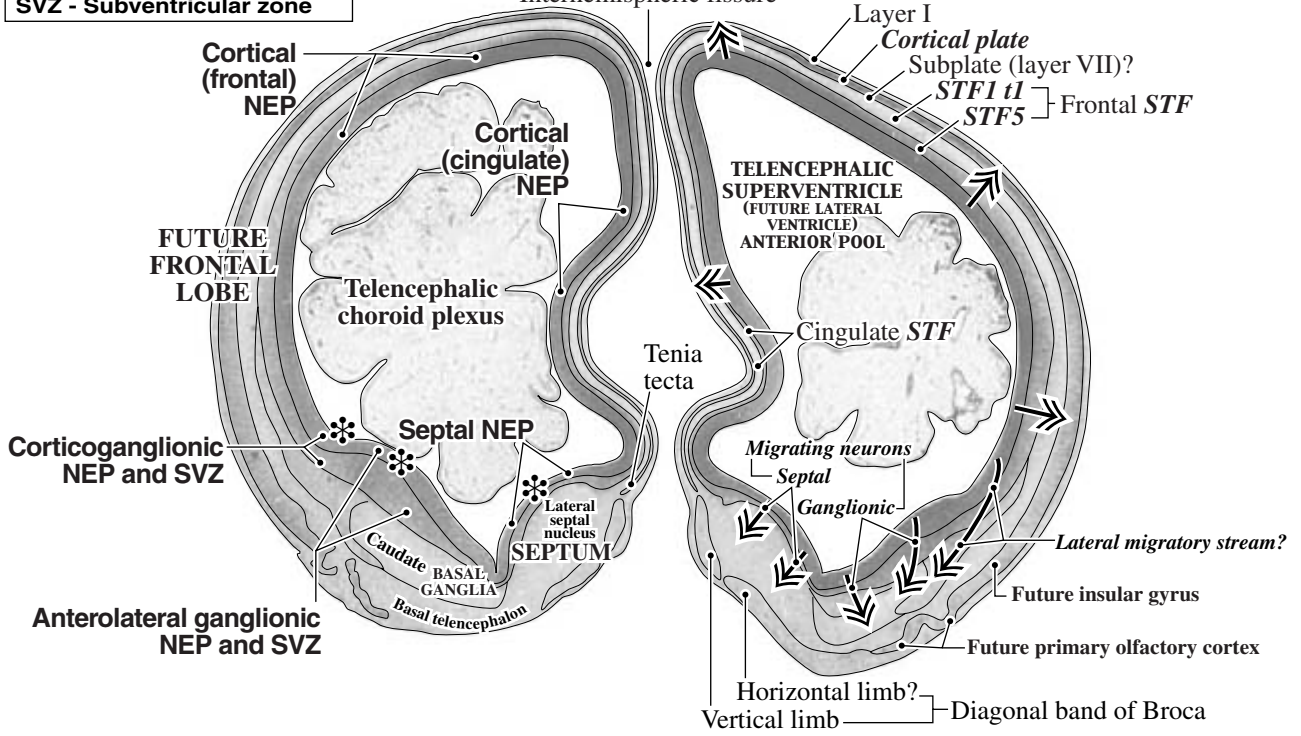
2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
NEP - Neuroepithelium
SVZ - Subventricular zone

Level 3: Section 440

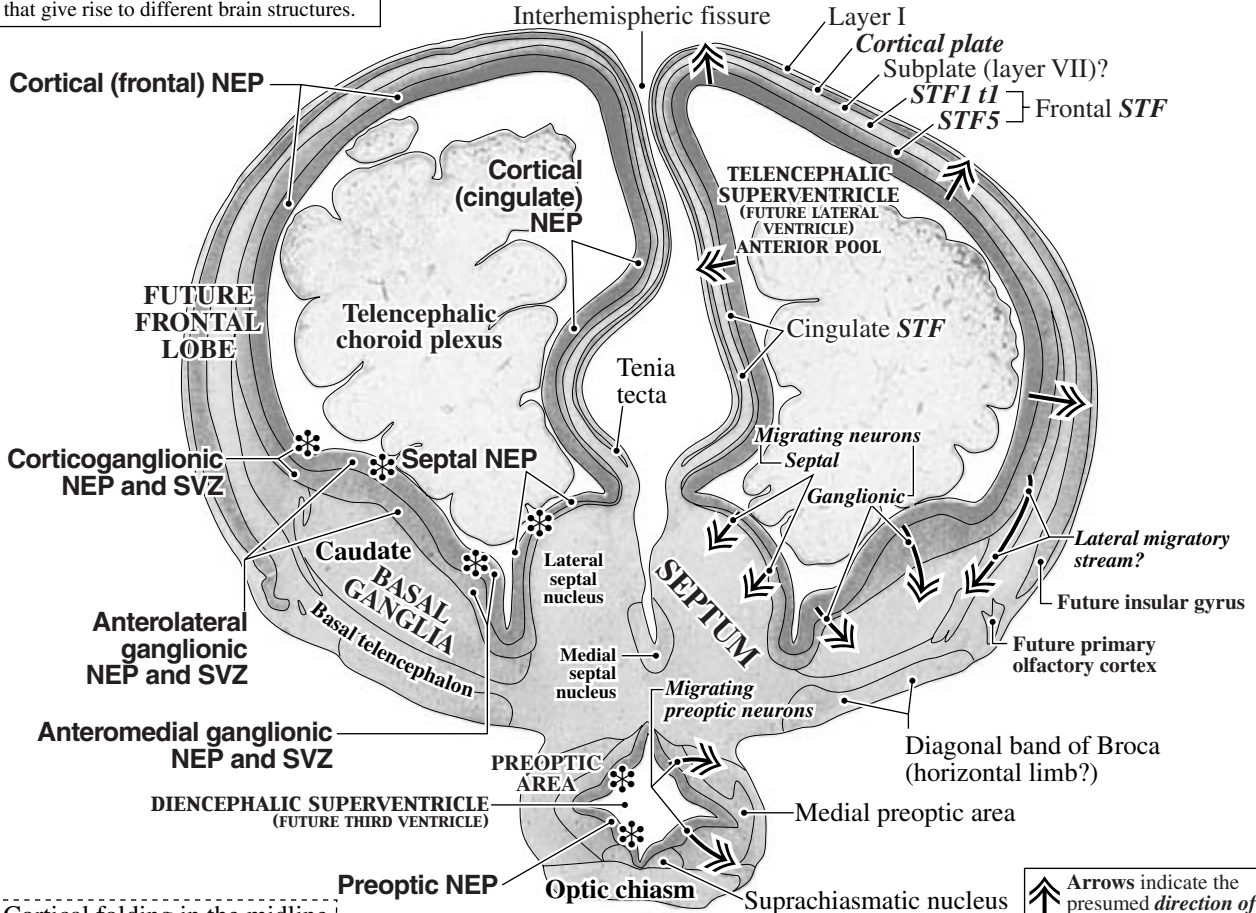
Interhemispheric fissure



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

Level 4: Section 410

Interhemispheric fissure

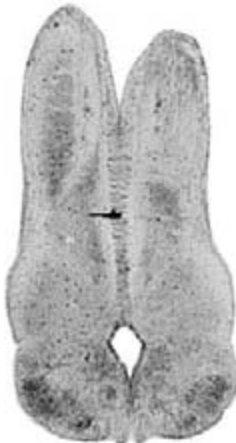
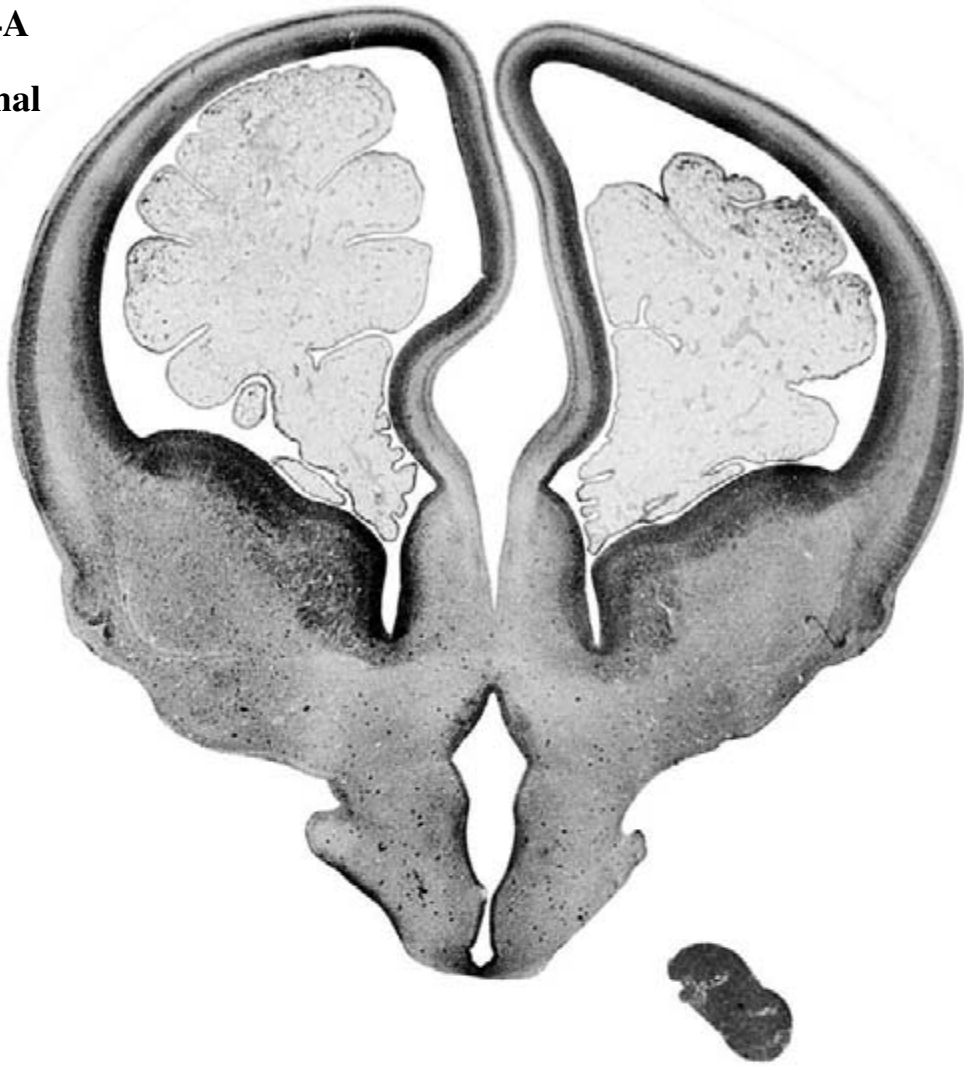


Cortical folding in the midline is a shrinkage artifact.

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 104A

GW9 Coronal
CR 42 mm
M841
Level 5:
Section 375



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:

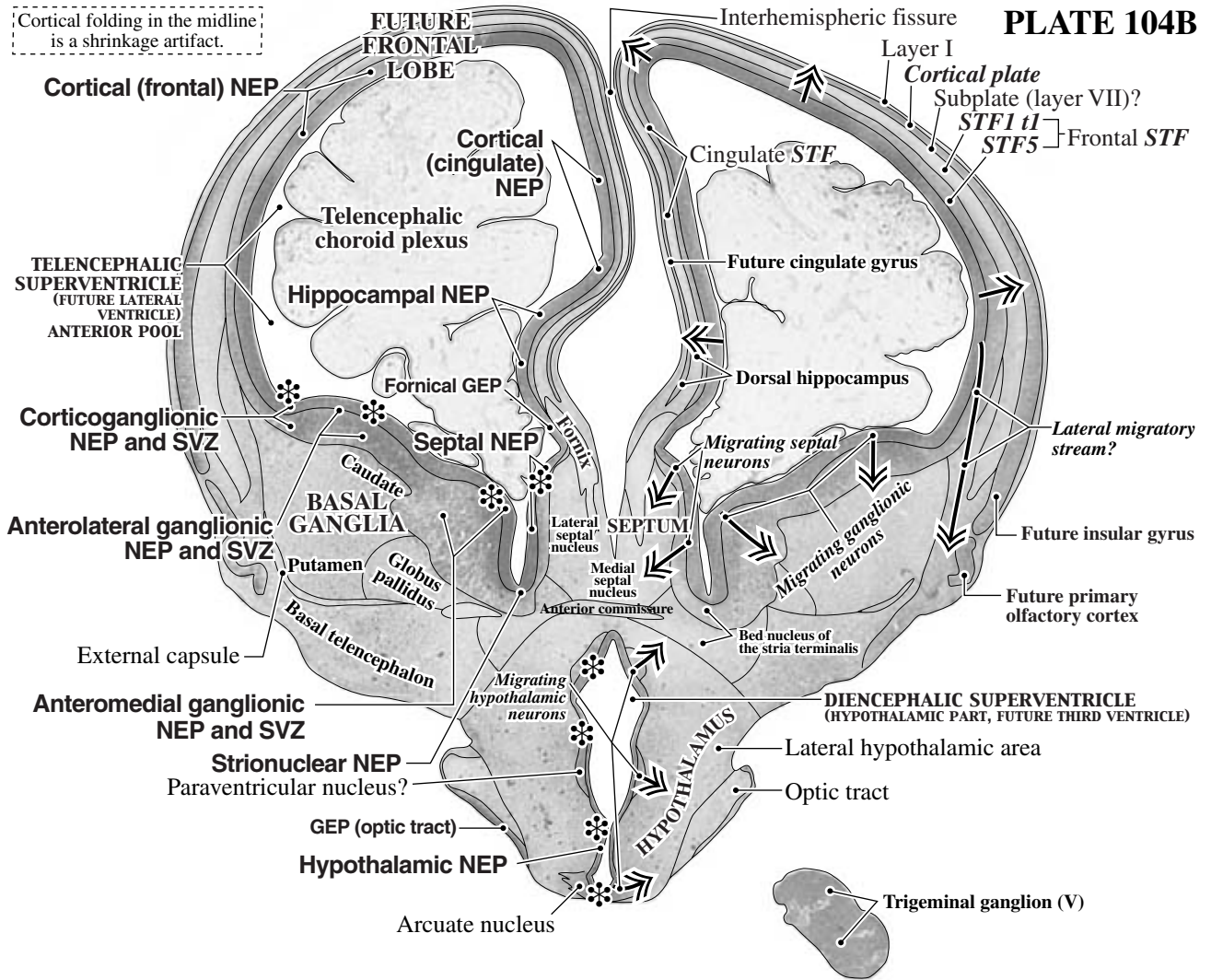
VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or Bold

Cortical folding in the midline is a shrinkage artifact.



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

SPINAL CORD

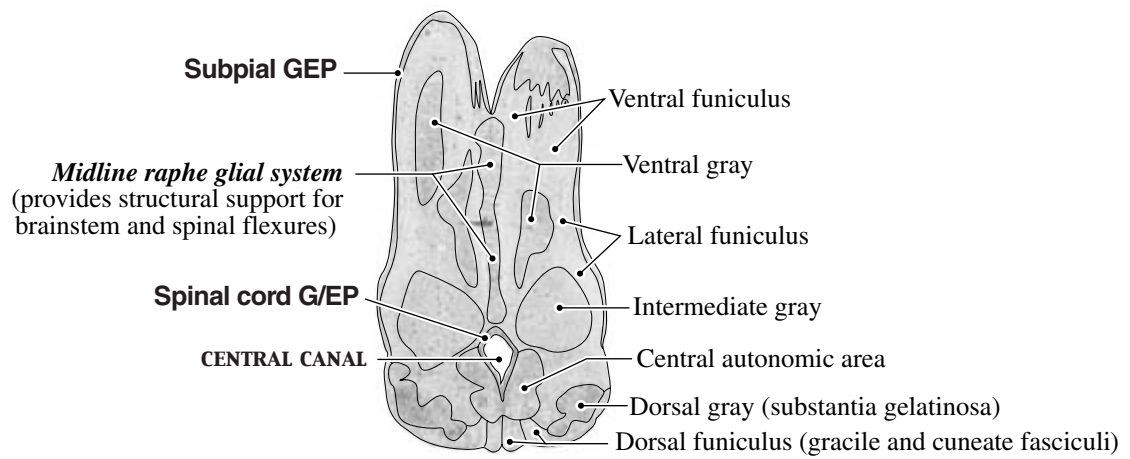
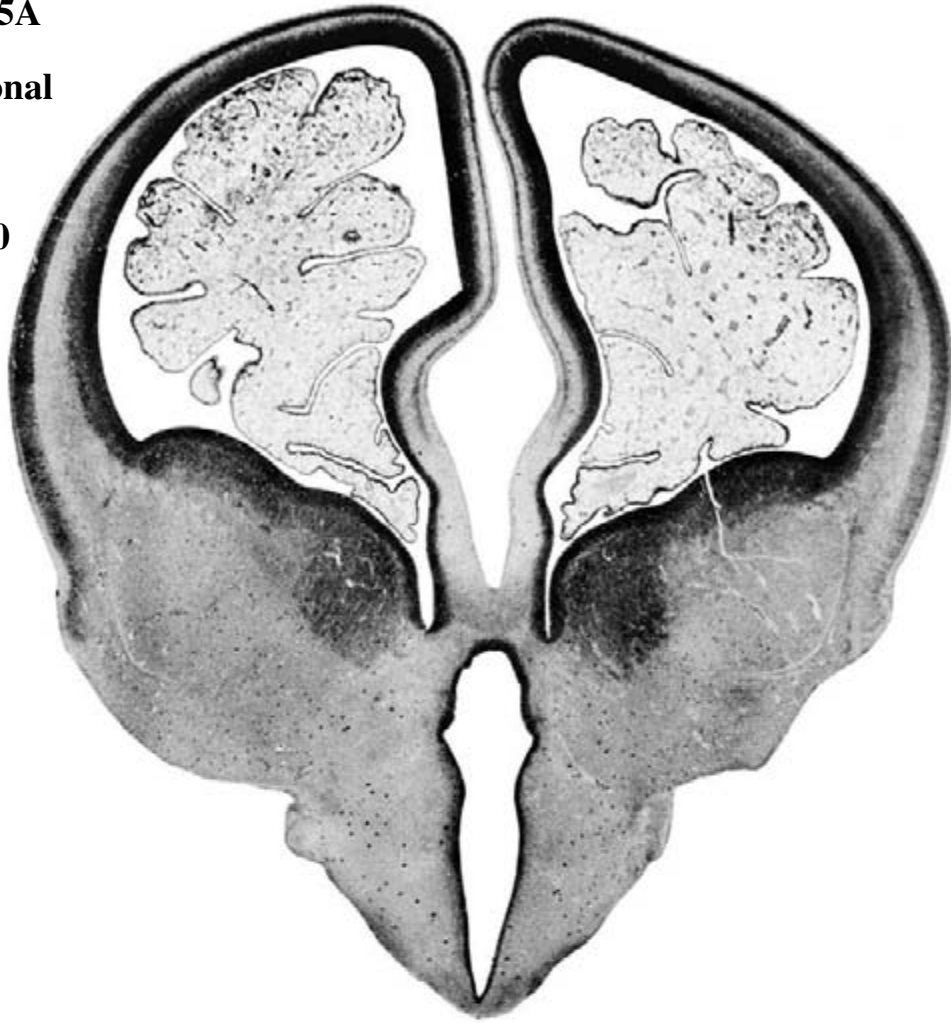


PLATE 105A

GW9 Coronal
CR 42 mm
M841
Level 6:
Section 360

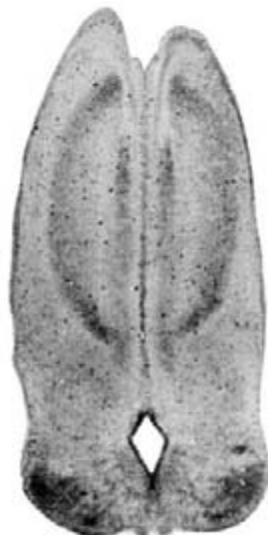


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

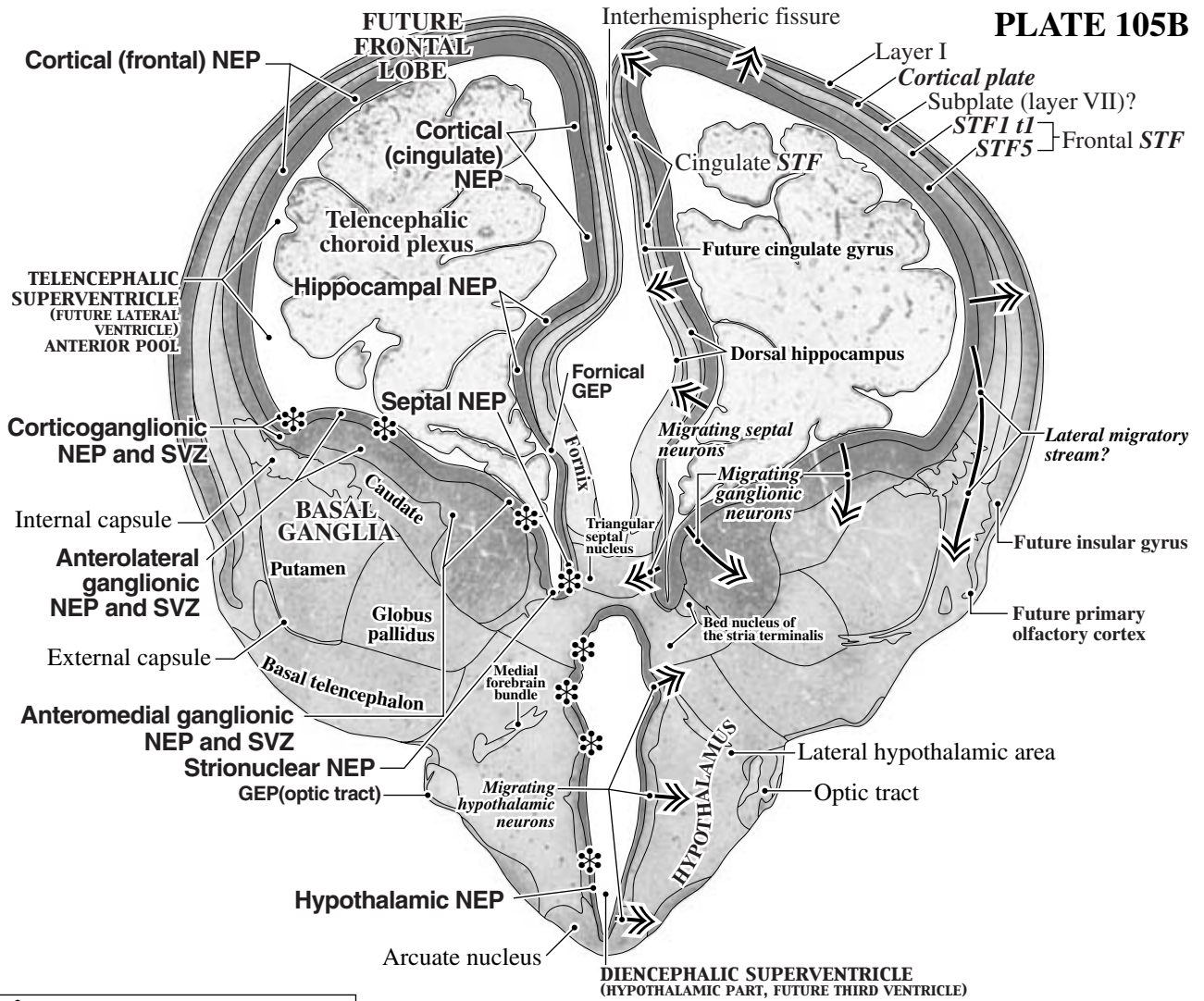
STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



2 mm

PLATE 105B

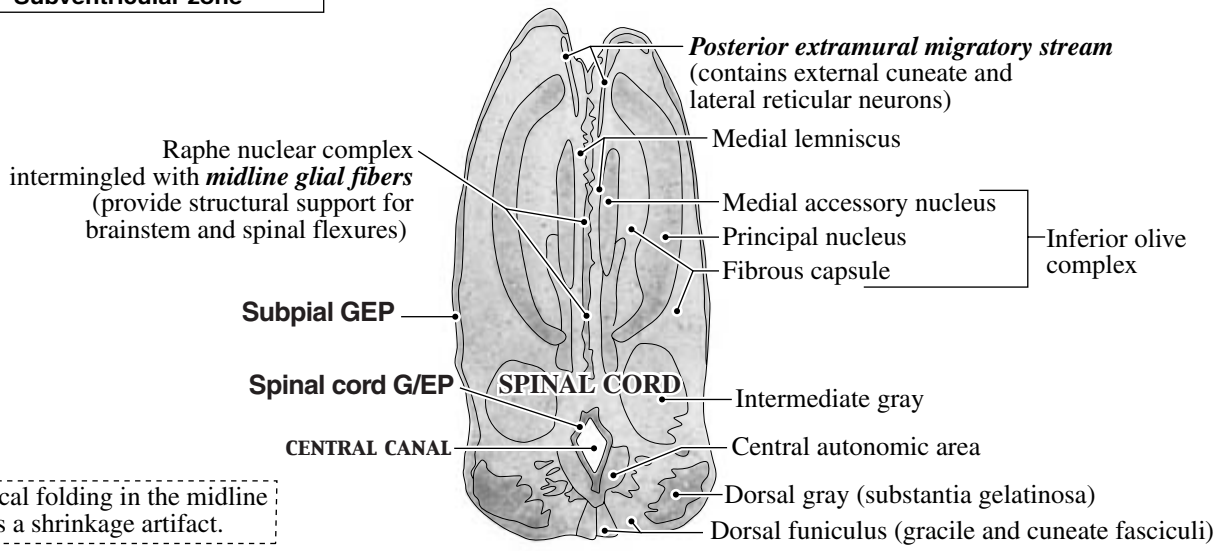


* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

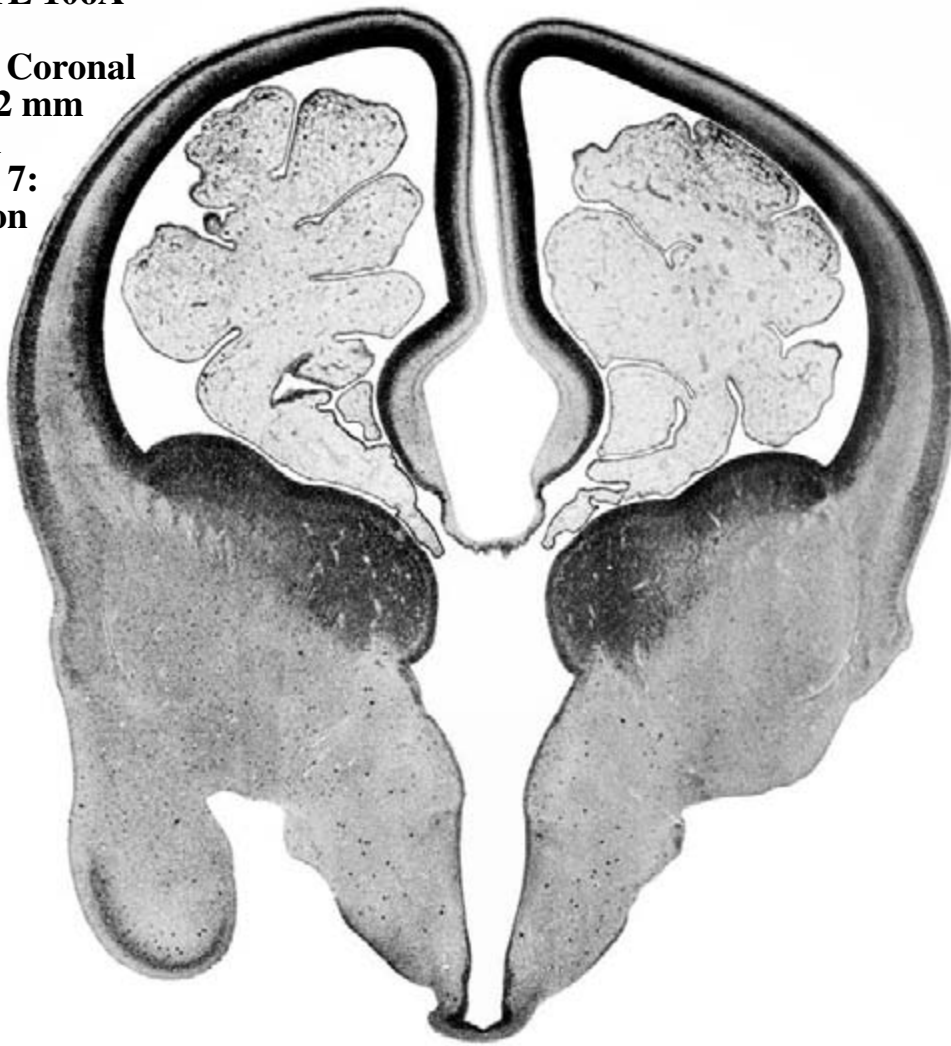
MEDULLA



Cortical folding in the midline is a shrinkage artifact.

PLATE 106A

GW9 Coronal
CR 42 mm
M841
Level 7:
Section
340



2 mm

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.
- STF4** Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:

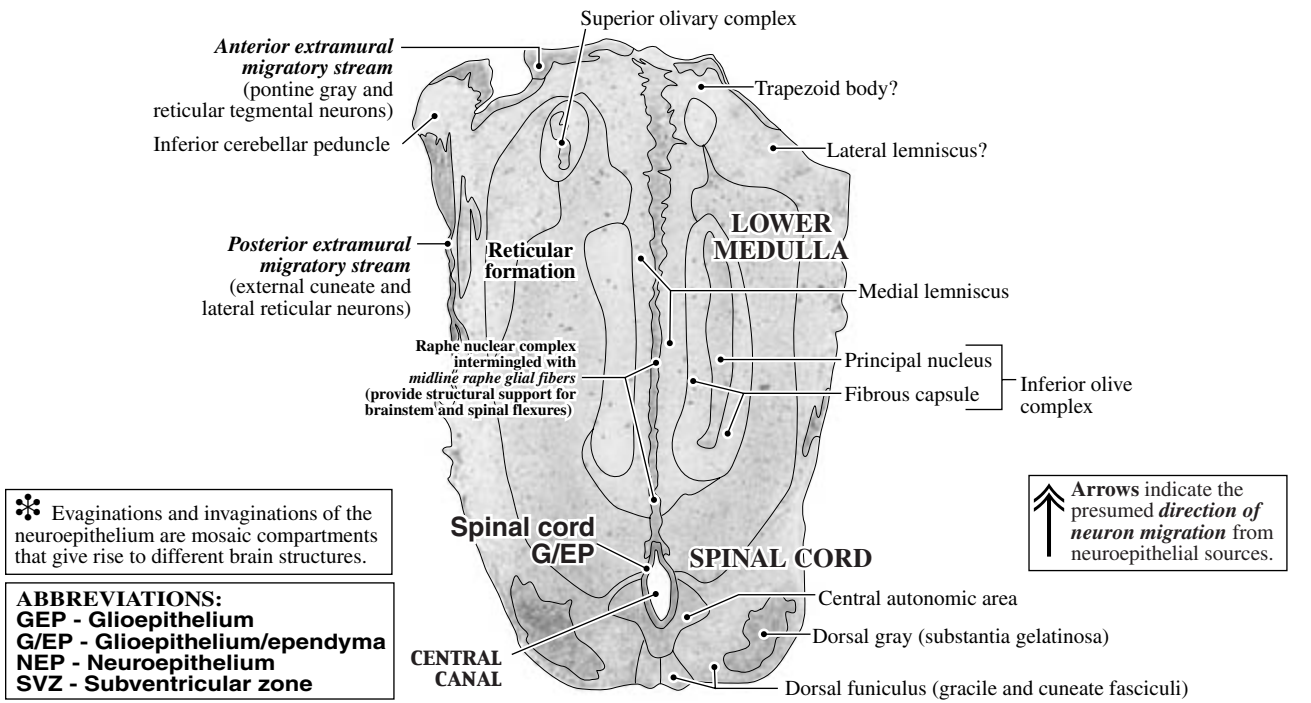
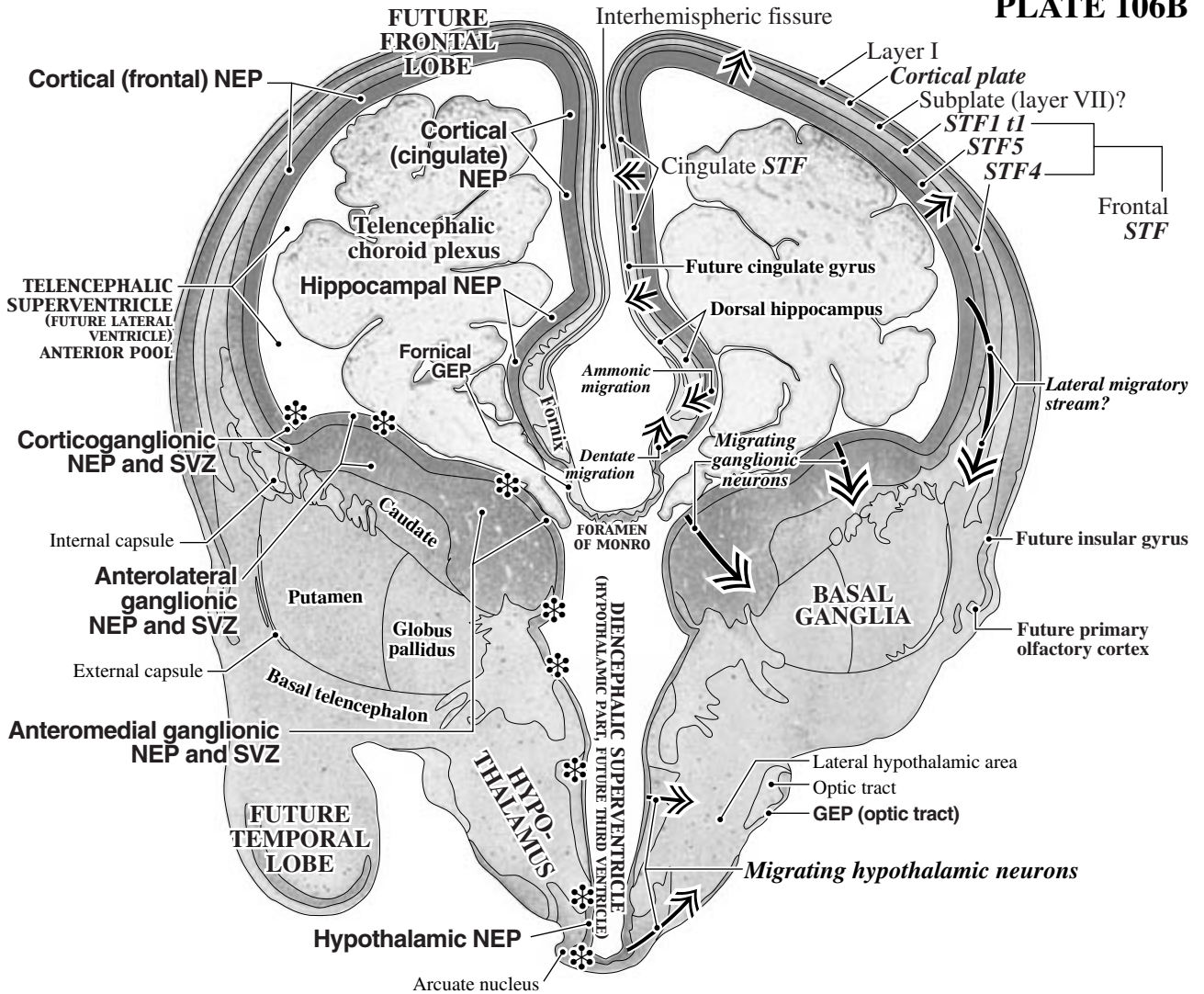
VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**

PLATE 106B



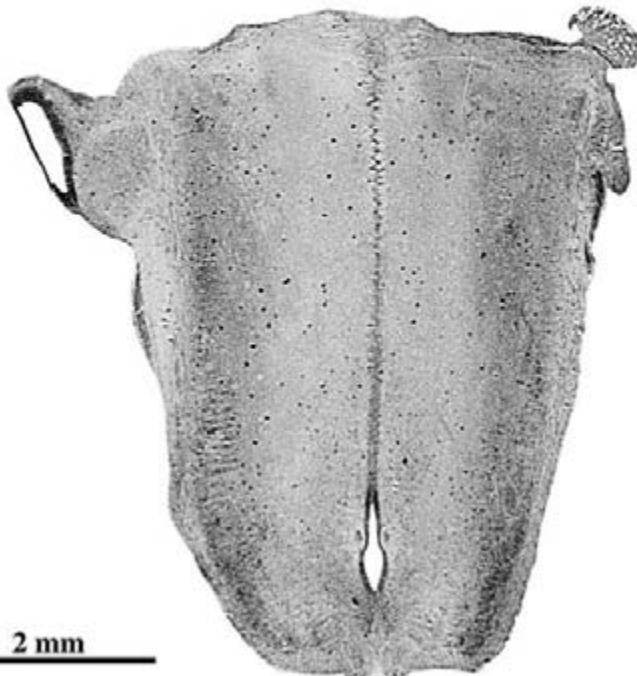
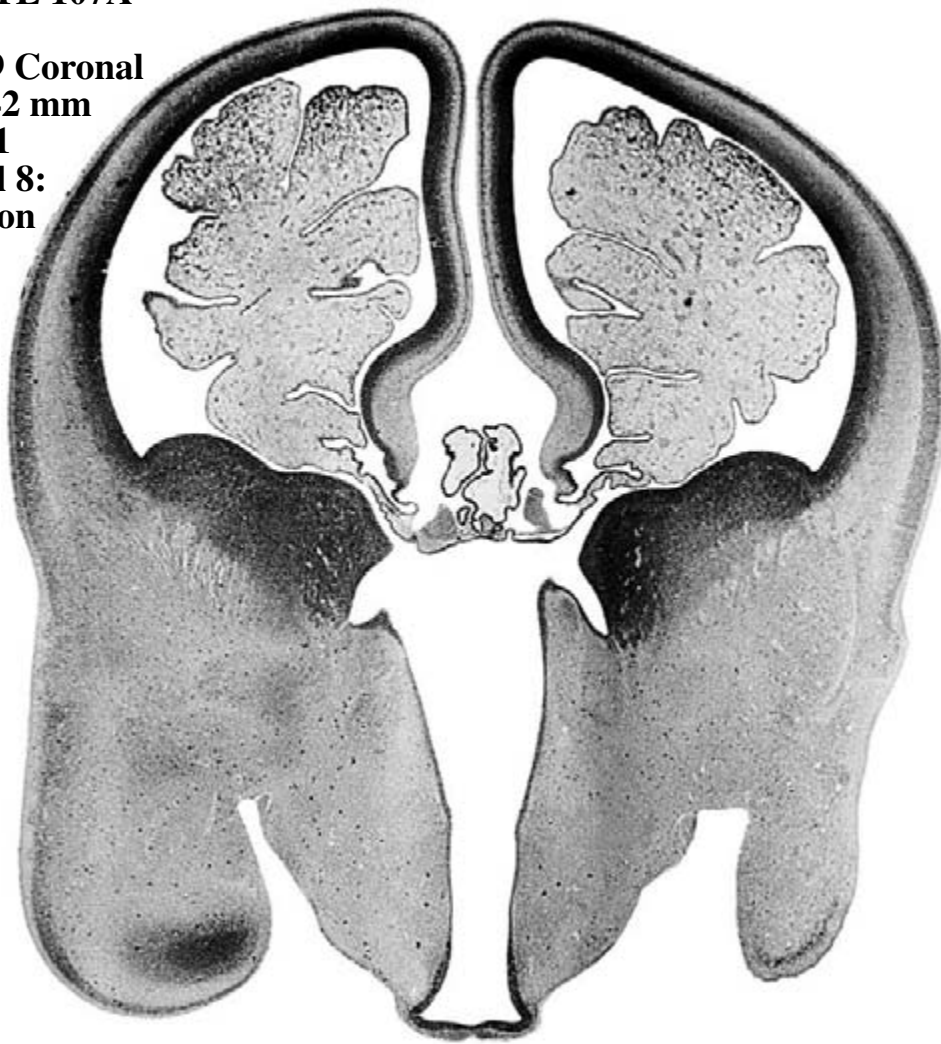
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
G/EP - Glioepithelium/ependyma
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 107A

GW9 Coronal
CR 42 mm
M841
Level 8:
Section
320



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

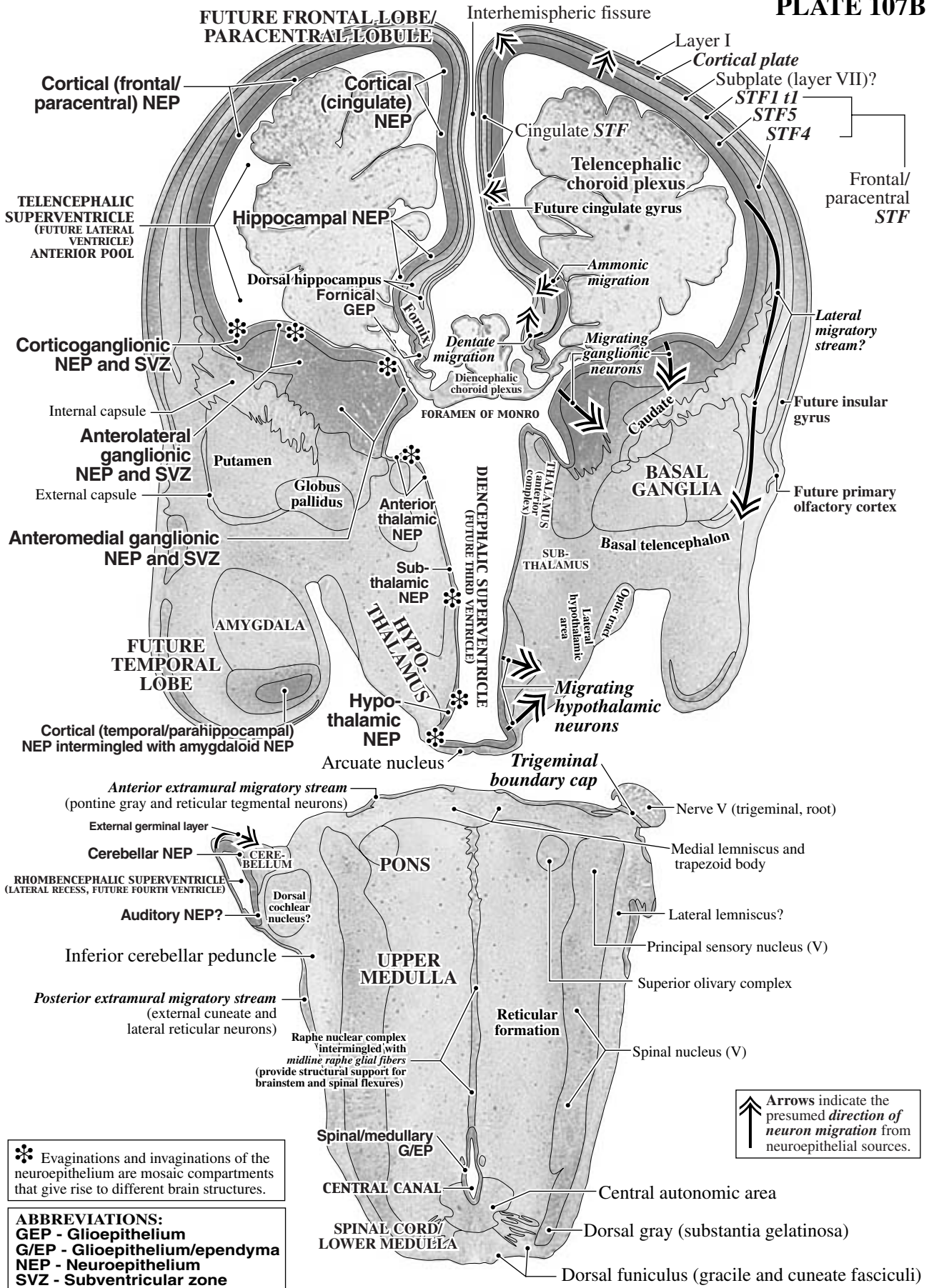
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**



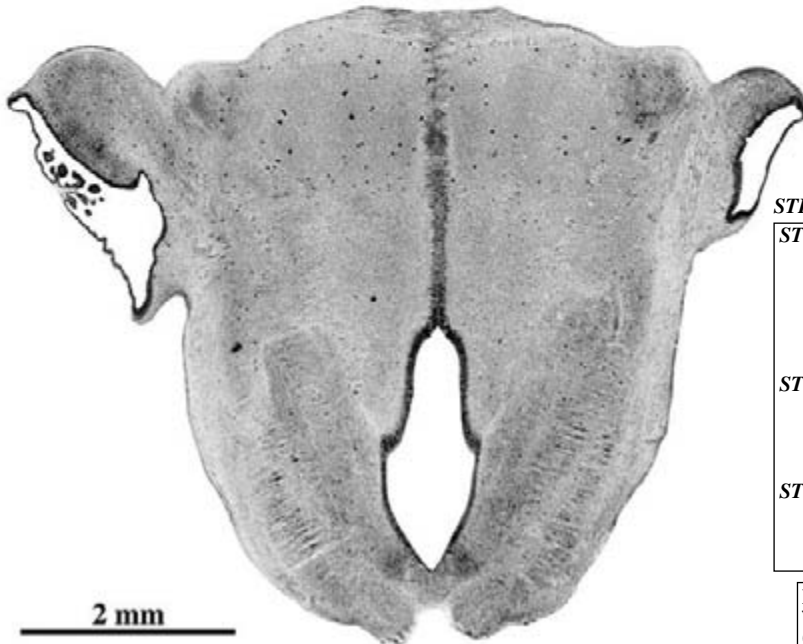
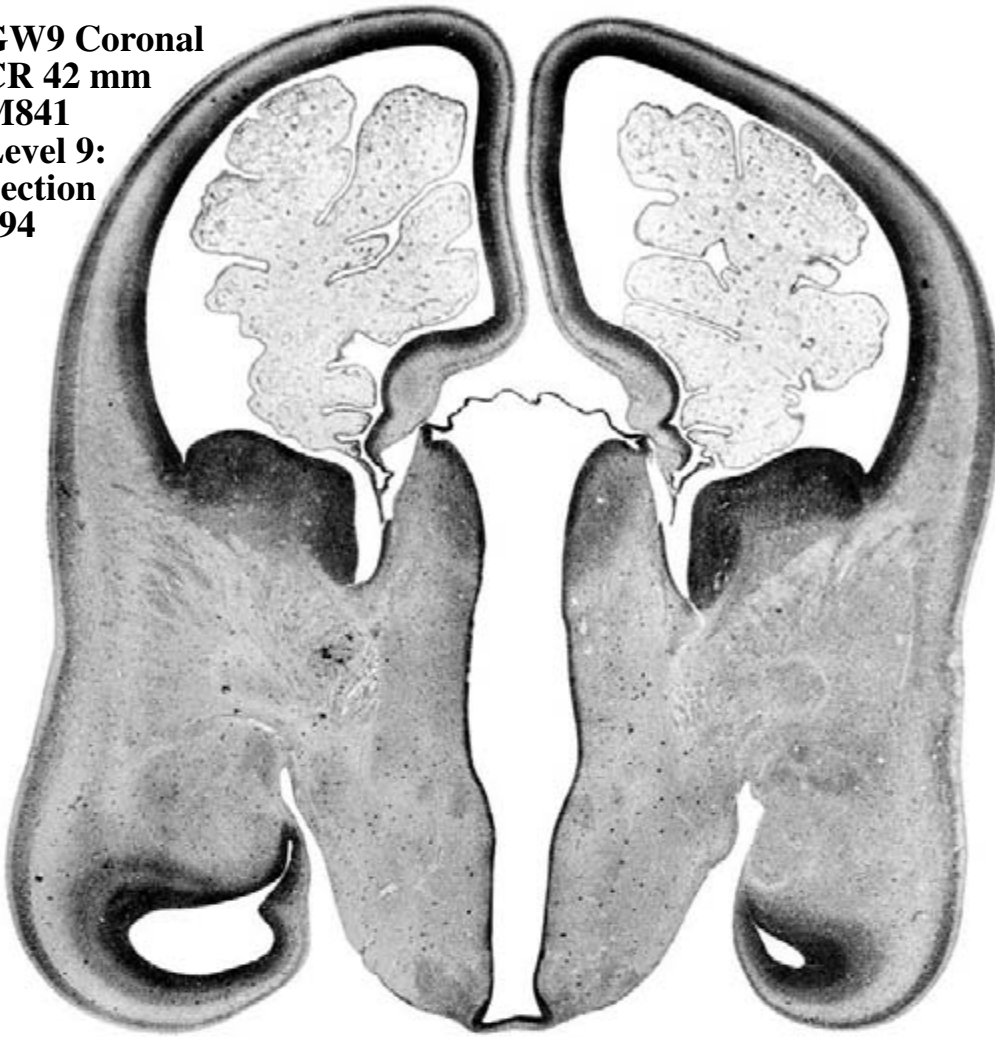
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
 GEP - Glioepithelium
 G/EP - Glioepithelium/ependyma
 NEP - Neuroepithelium
 SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 108A

GW9 Coronal
CR 42 mm
M841
Level 9:
Section
294



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

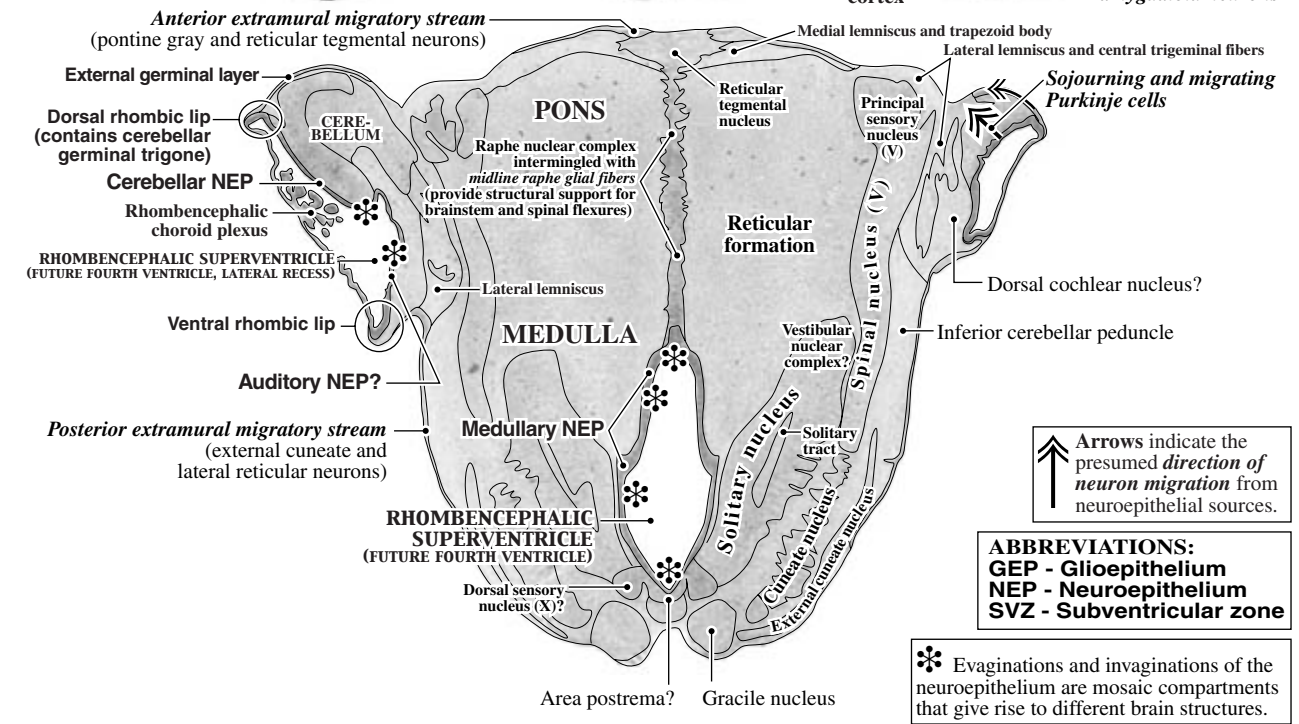
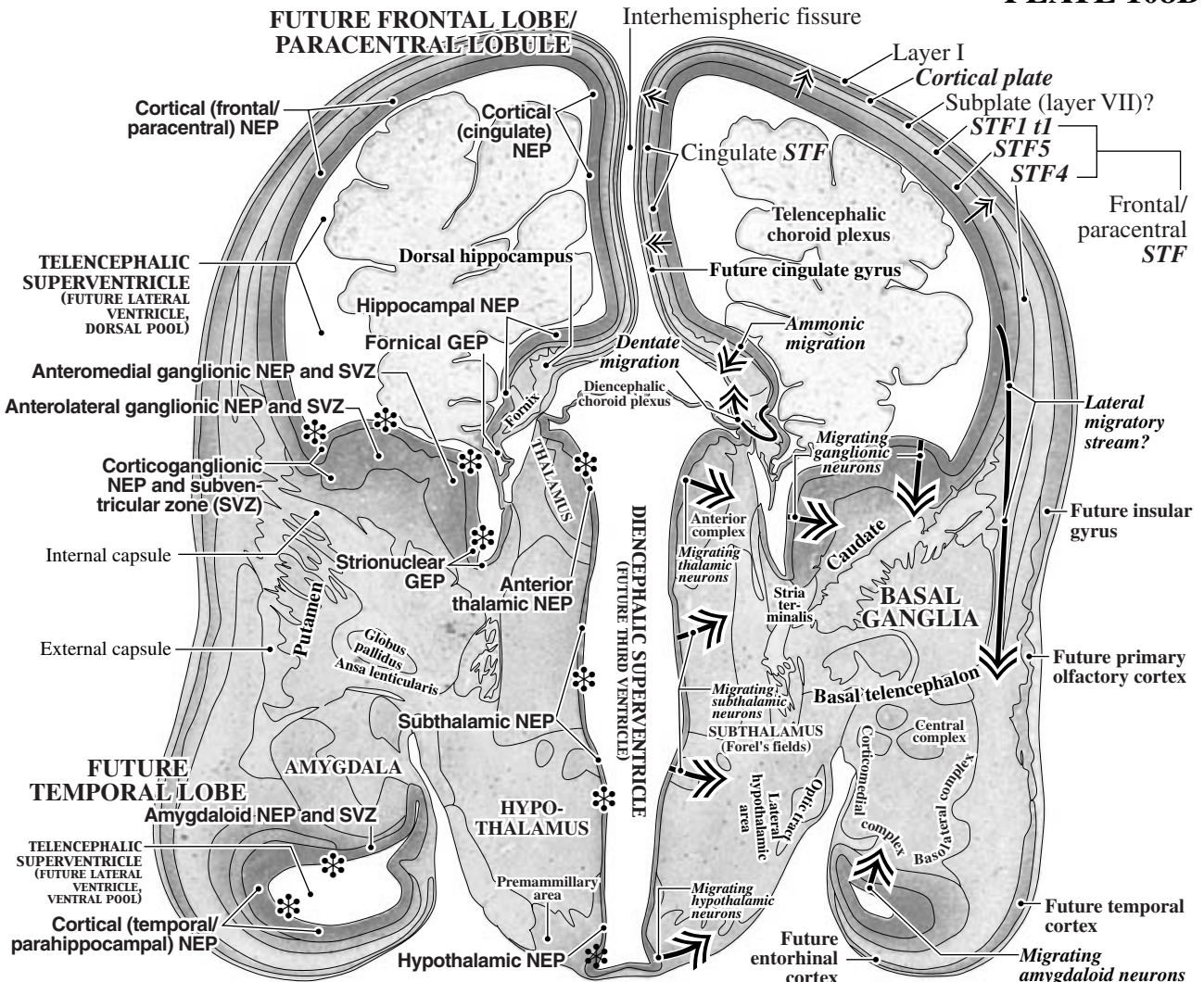
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**



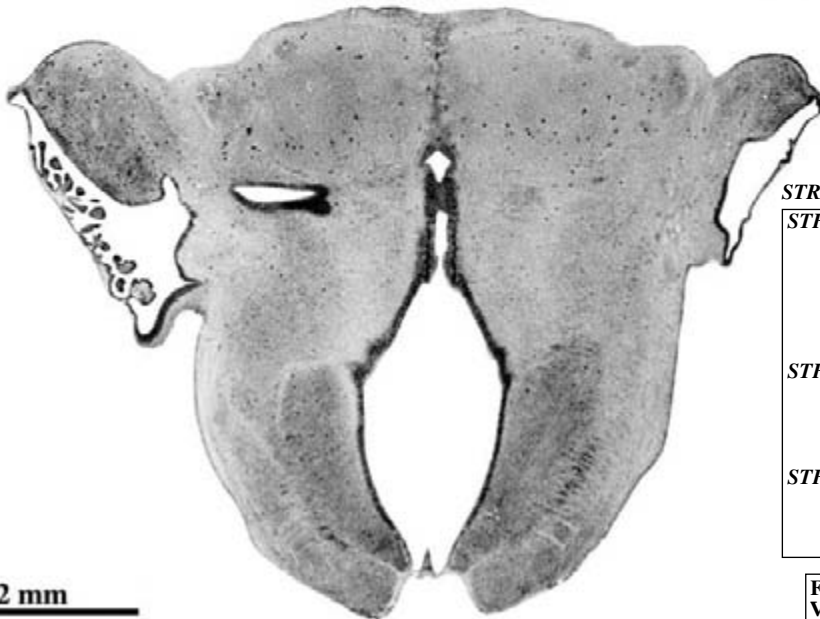
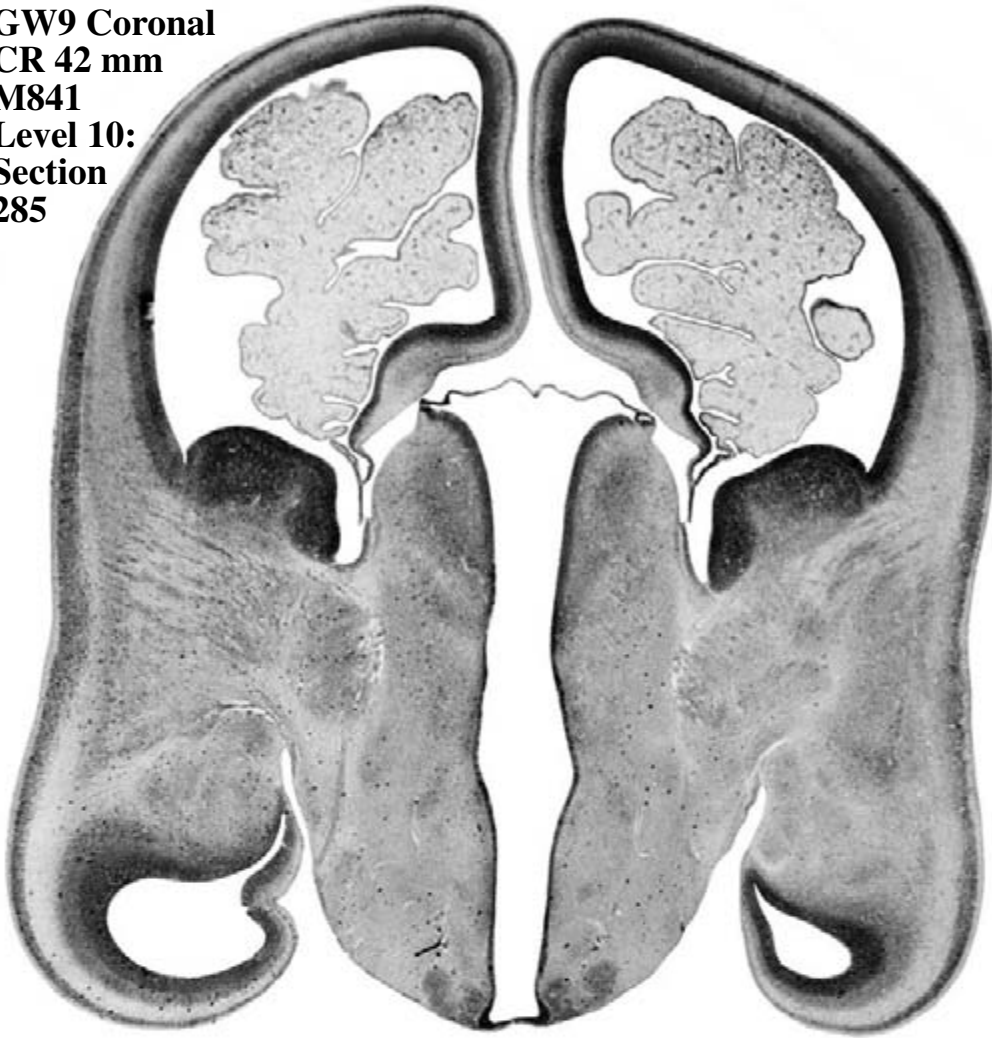
↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioeptithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 109A

GW9 Coronal
CR 42 mm
M841
Level 10:
Section
285



2 mm

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**

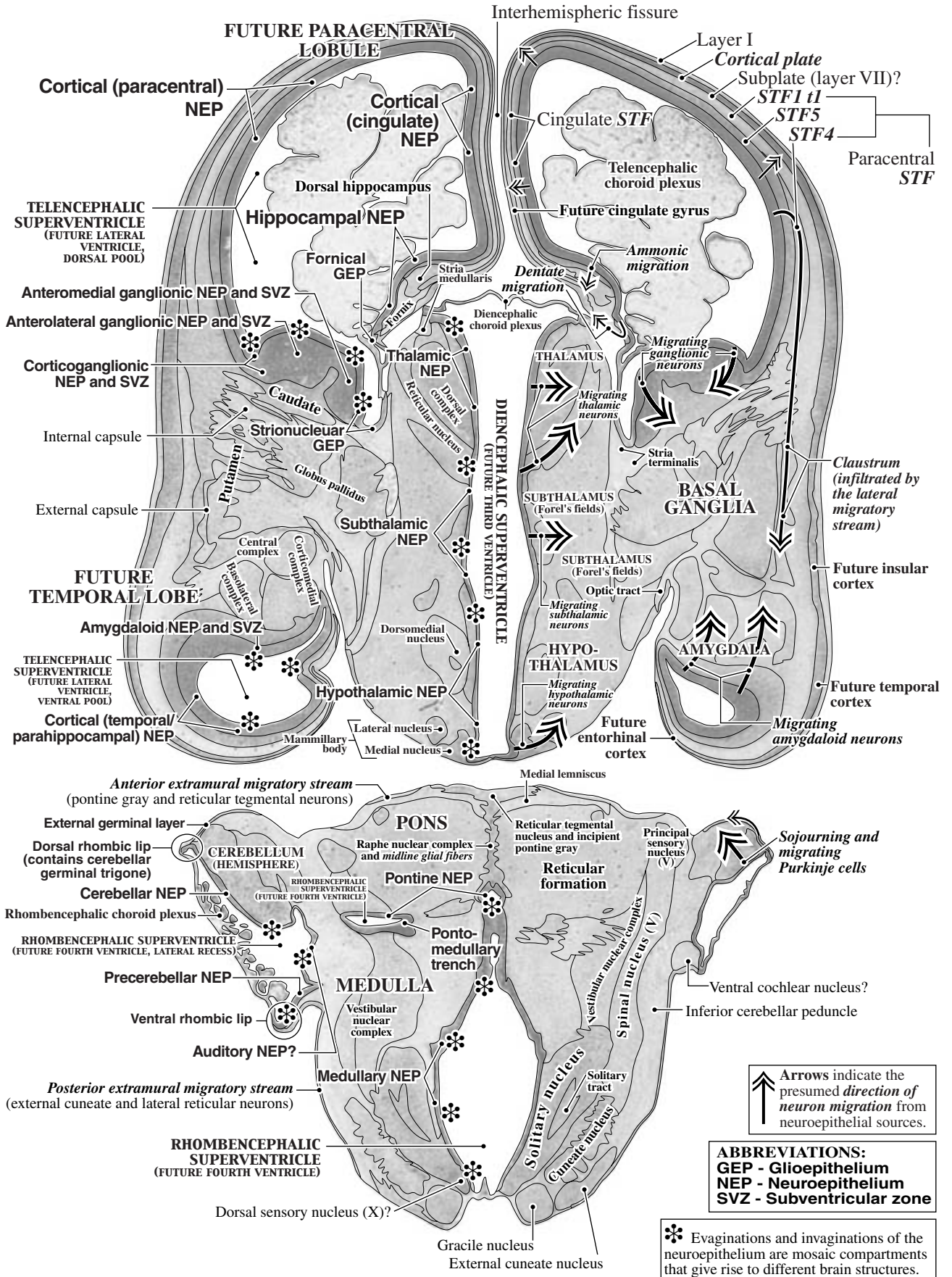
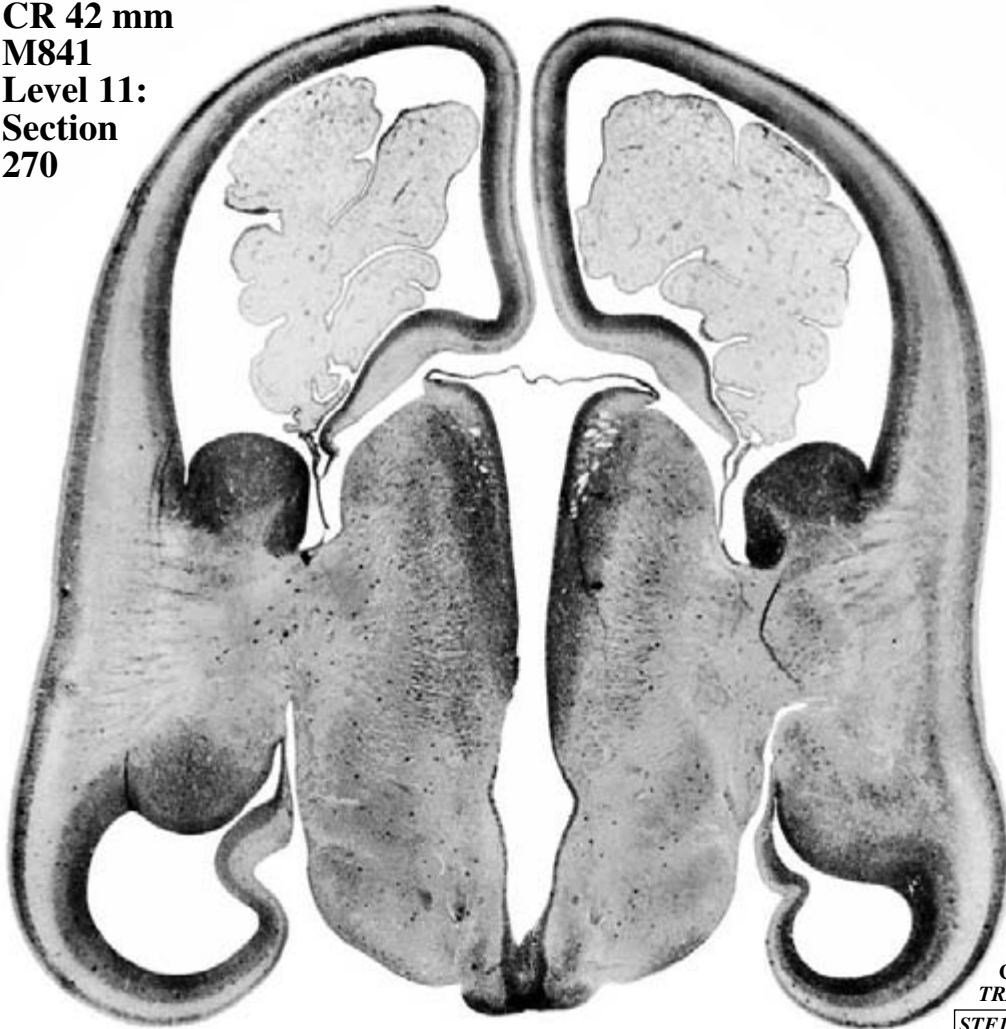


PLATE 110A

GW9 Coronal
CR 42 mm
M841
Level 11:
Section
270

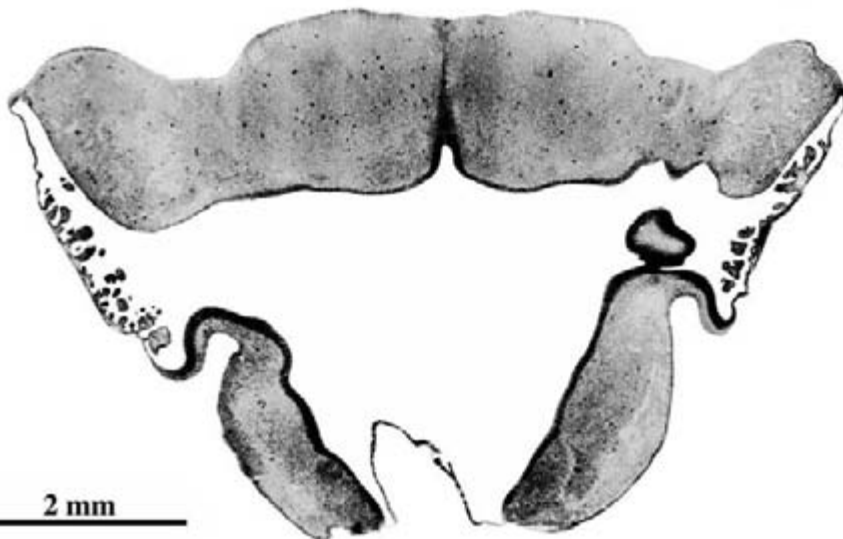


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

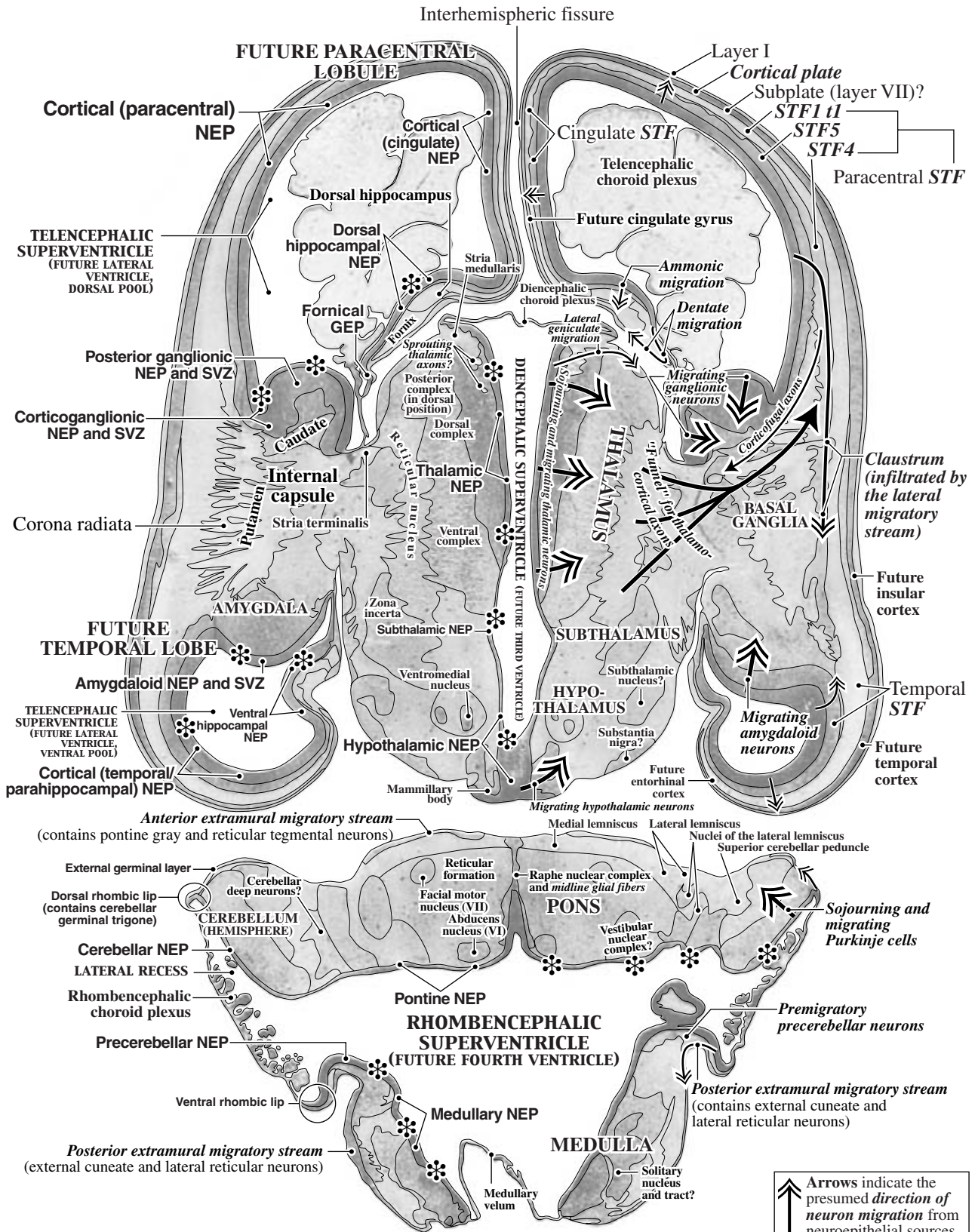
STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

PLATE 111A

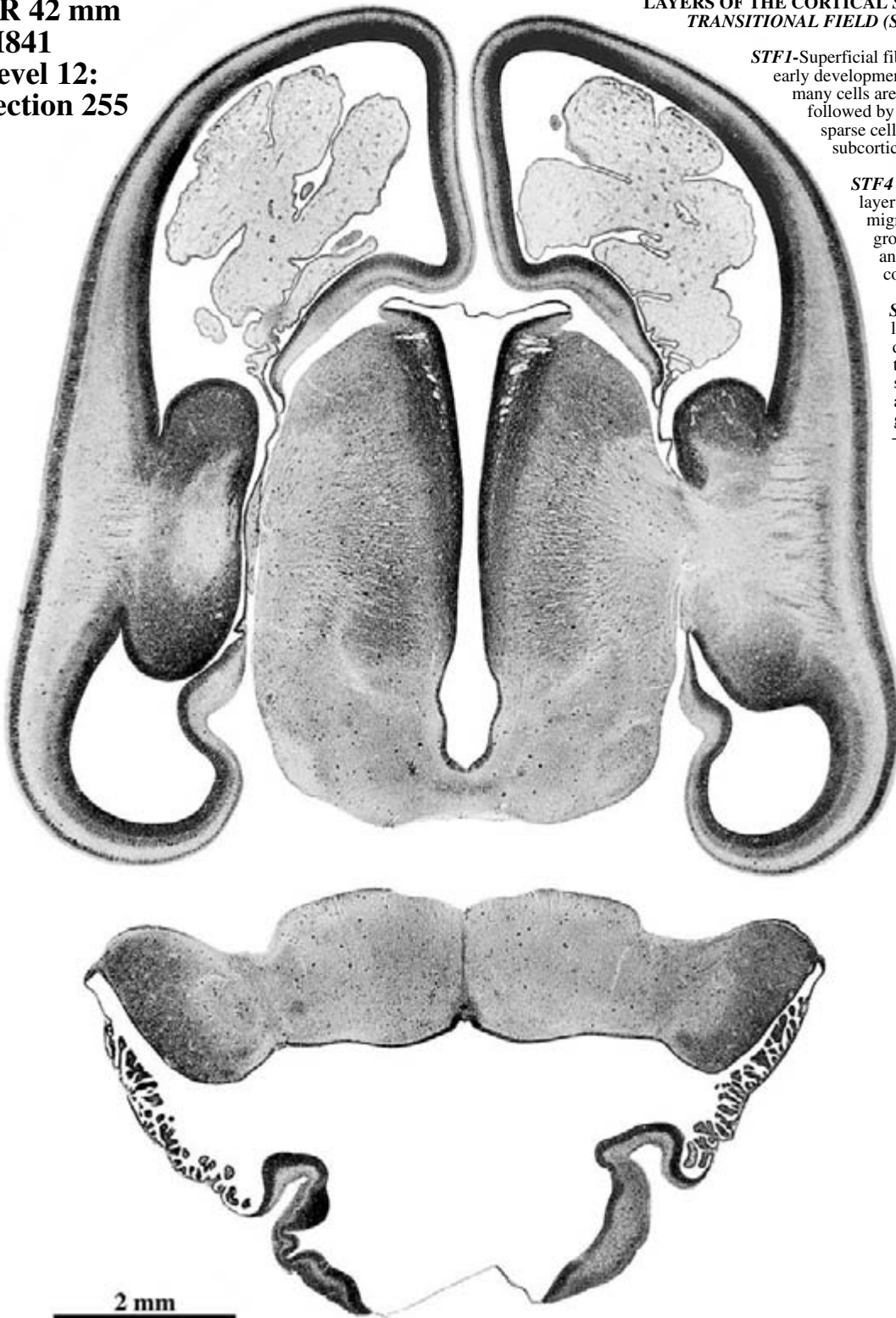
GW9 Coronal
CR 42 mm
M841
Level 12:
Section 255

LAYERS OF THE CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)

STF1-Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

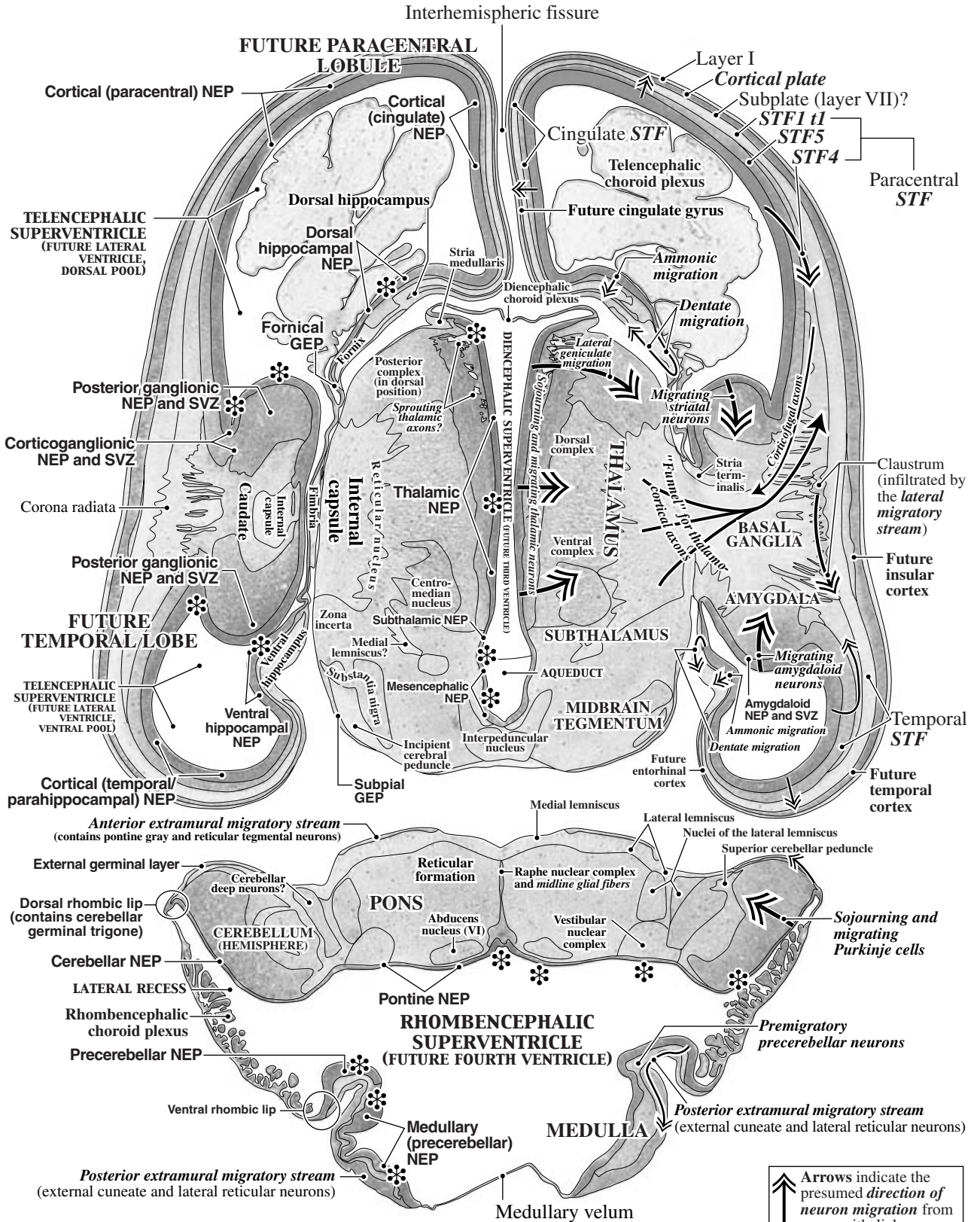
STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

↑ Arrows indicate the presumed direction of axon growth in brain fiber tracts.

PLATE 112A

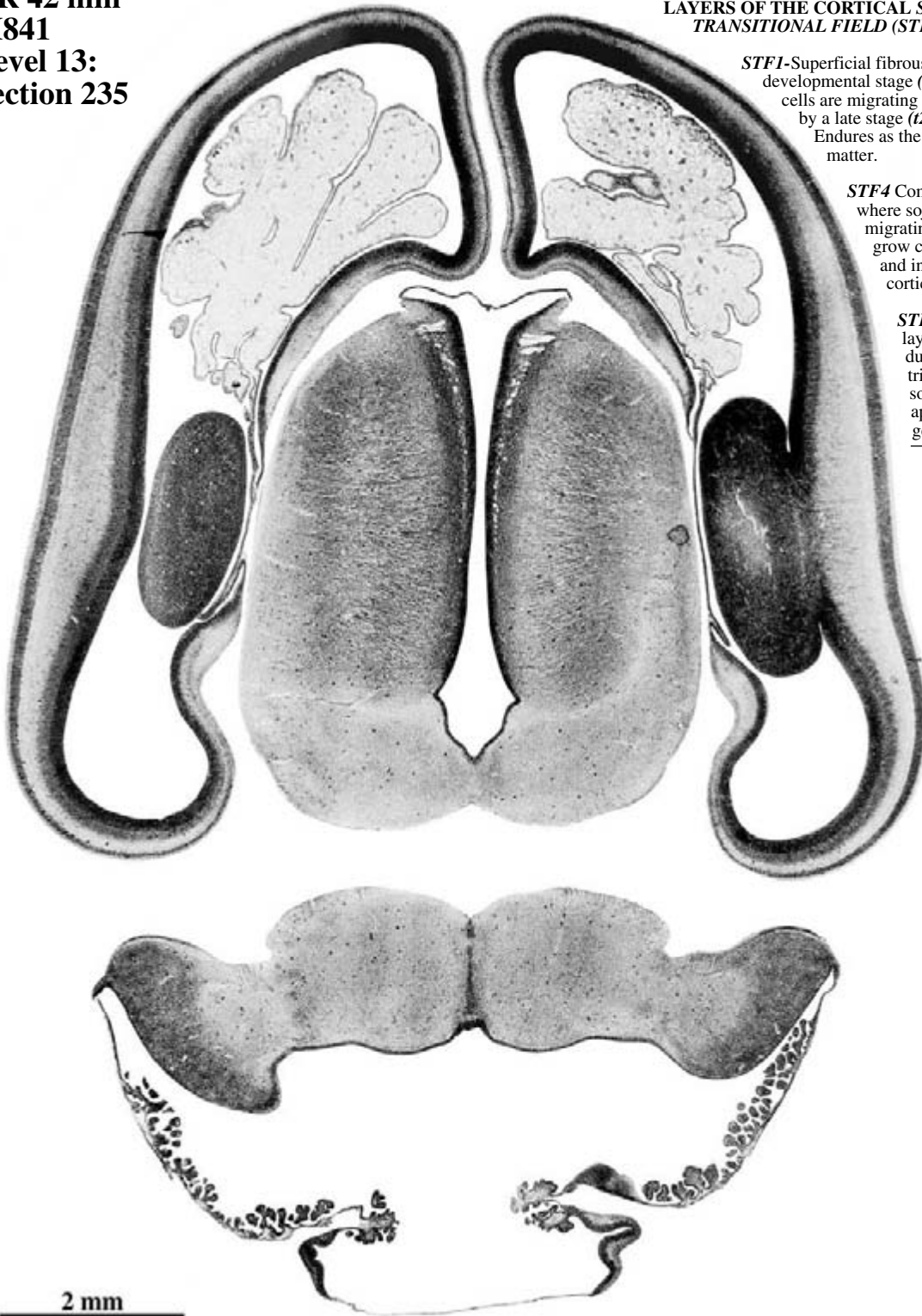
GW9 Coronal
CR 42 mm
M841
Level 13:
Section 235

LAYERS OF THE CORTICAL STRATIFIED
TRANSITIONAL FIELD (*STF*)

STF1-Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

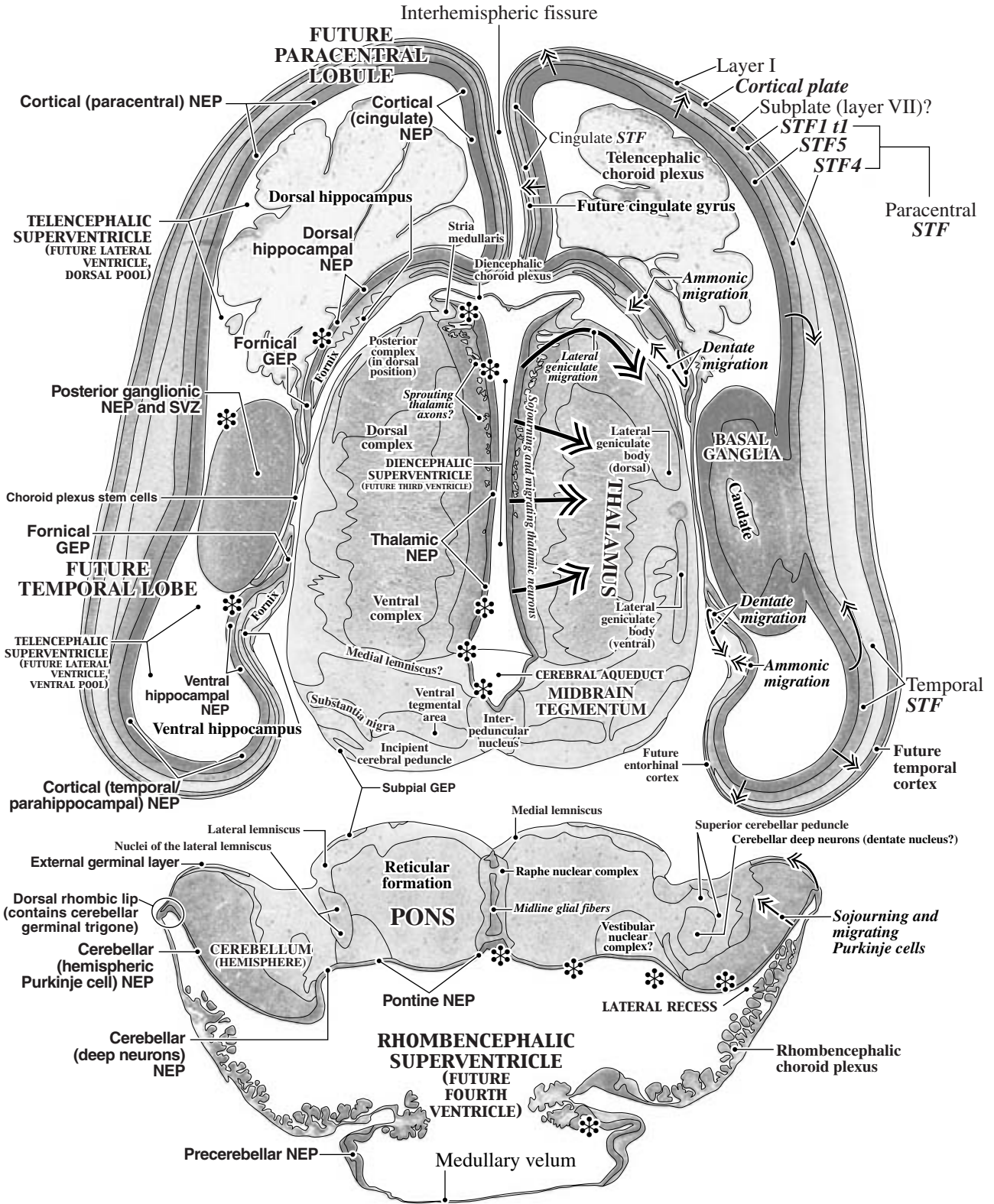
STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 113A

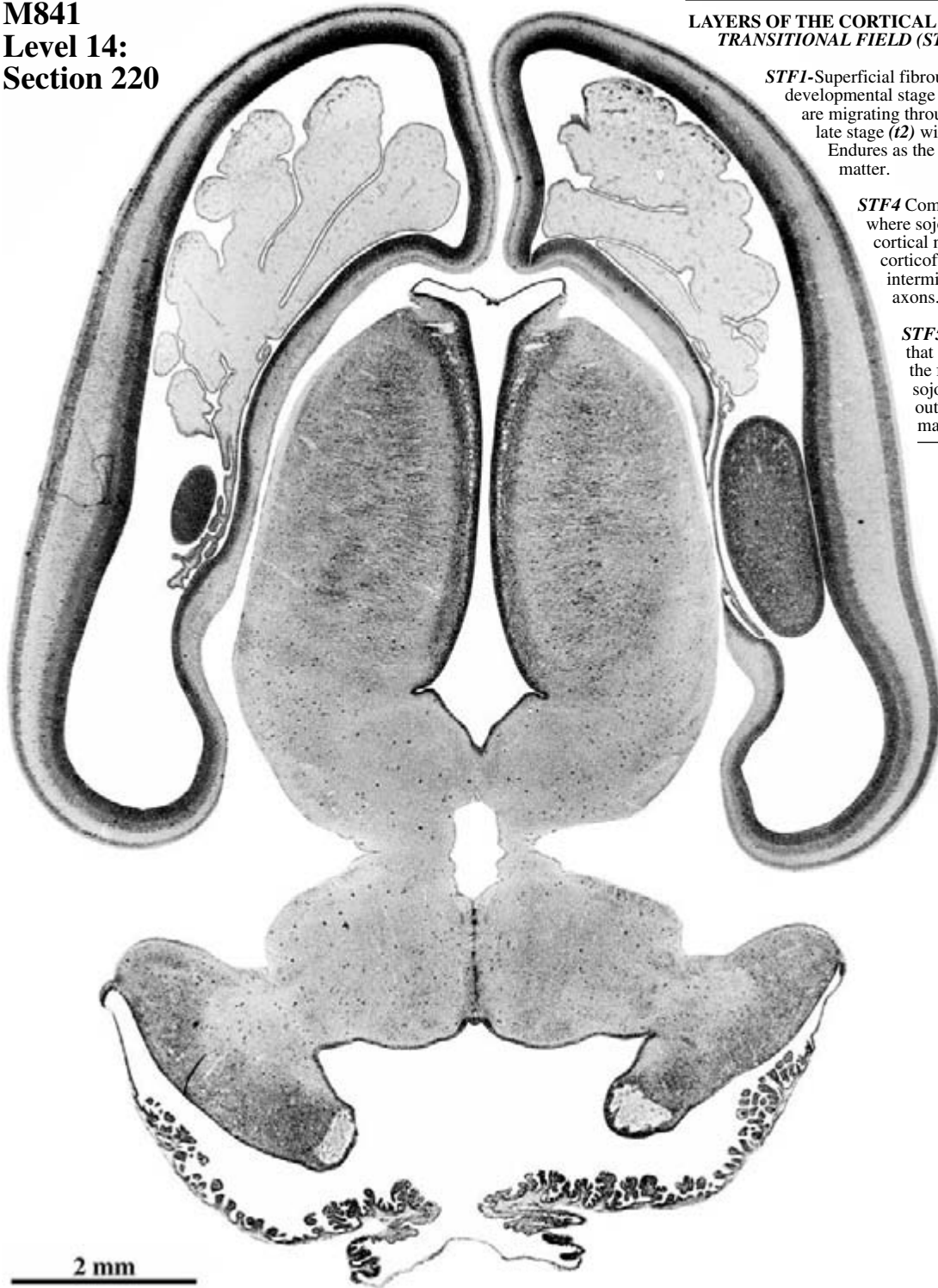
GW9 Coronal

CR 42 mm

M841

Level 14:

Section 220


**LAYERS OF THE CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

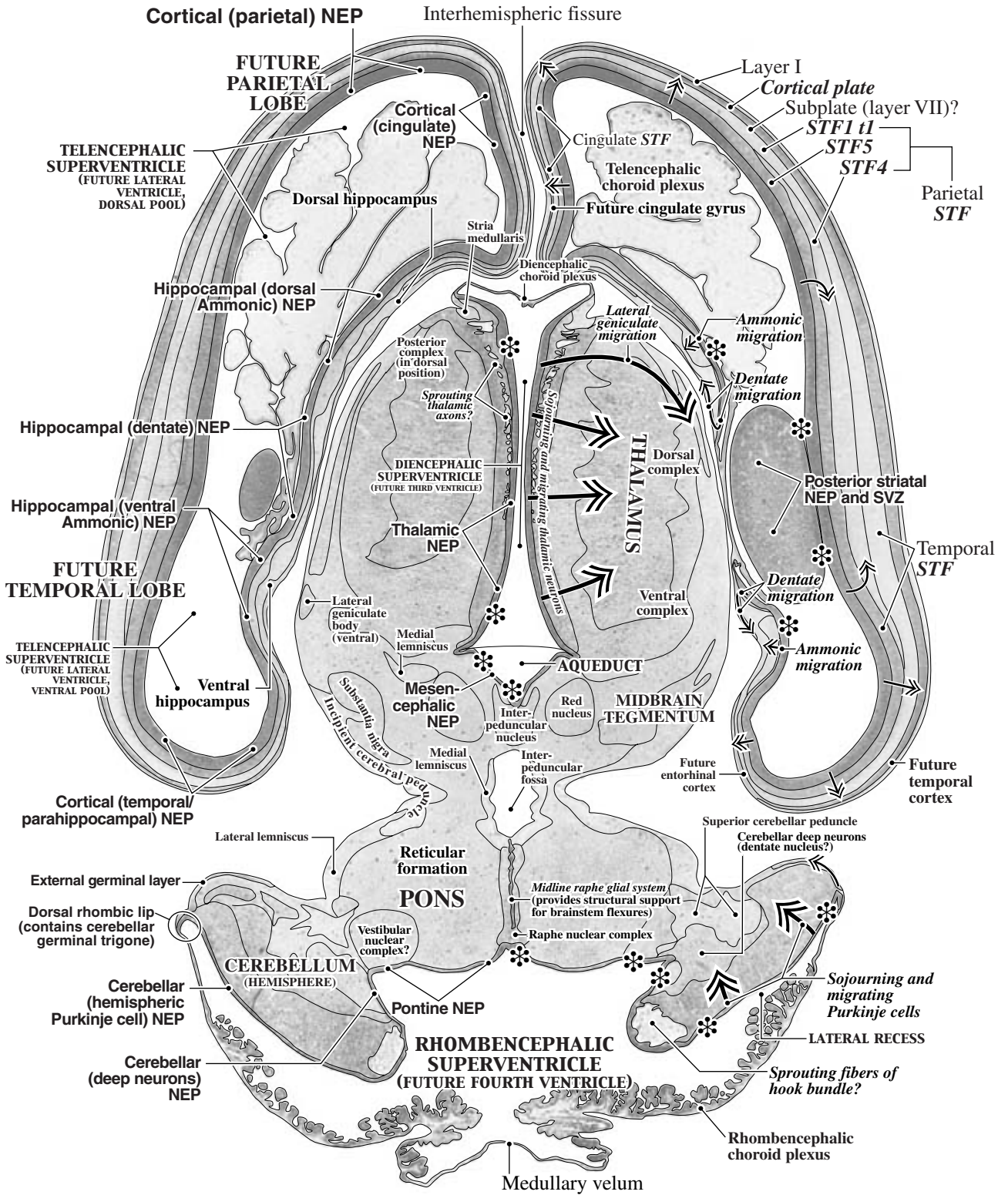
STF1-Superficial fibrous layer with an early developmental stage (***1I***) when many cells are migrating through it, followed by a late stage (***12***) with sparse cells. Endures as the subcortical white matter.

STF4 Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
 NEP - Neuroepithelium
 SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 114A

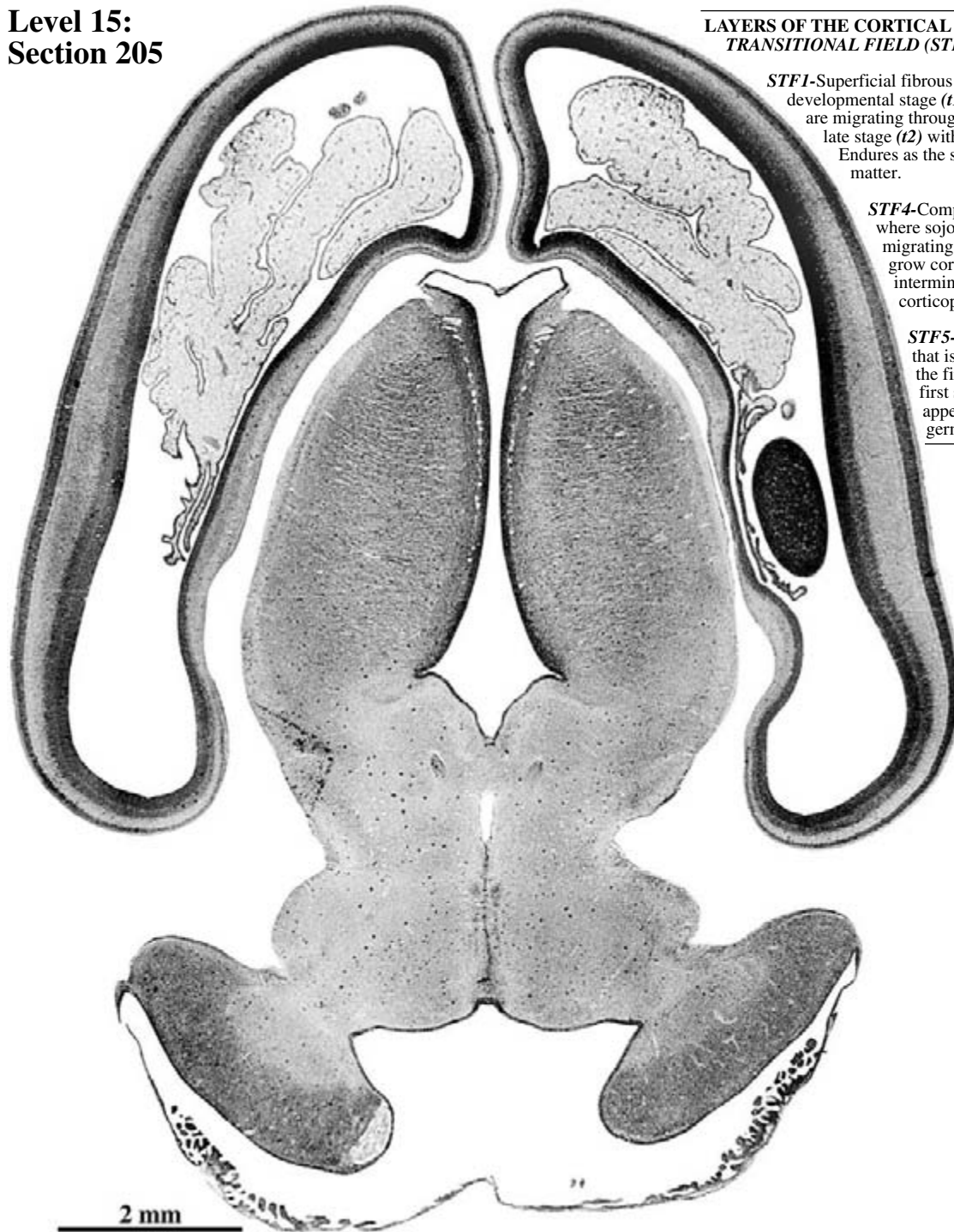
GW9 Coronal

CR 42 mm

M841

Level 15:

Section 205


**LAYERS OF THE CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

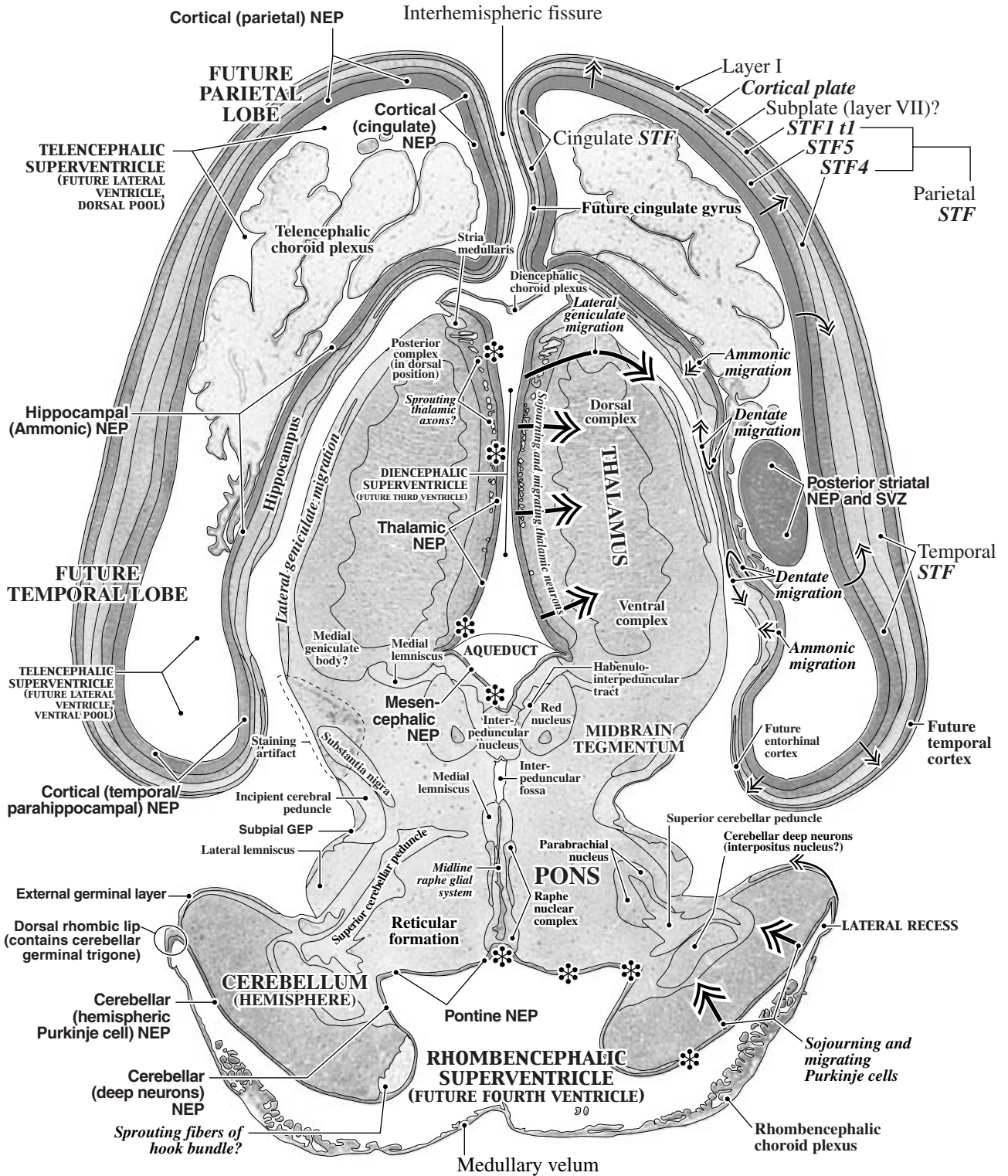
STF1-Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF4-Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 SVZ - Subventricular zone

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 115A

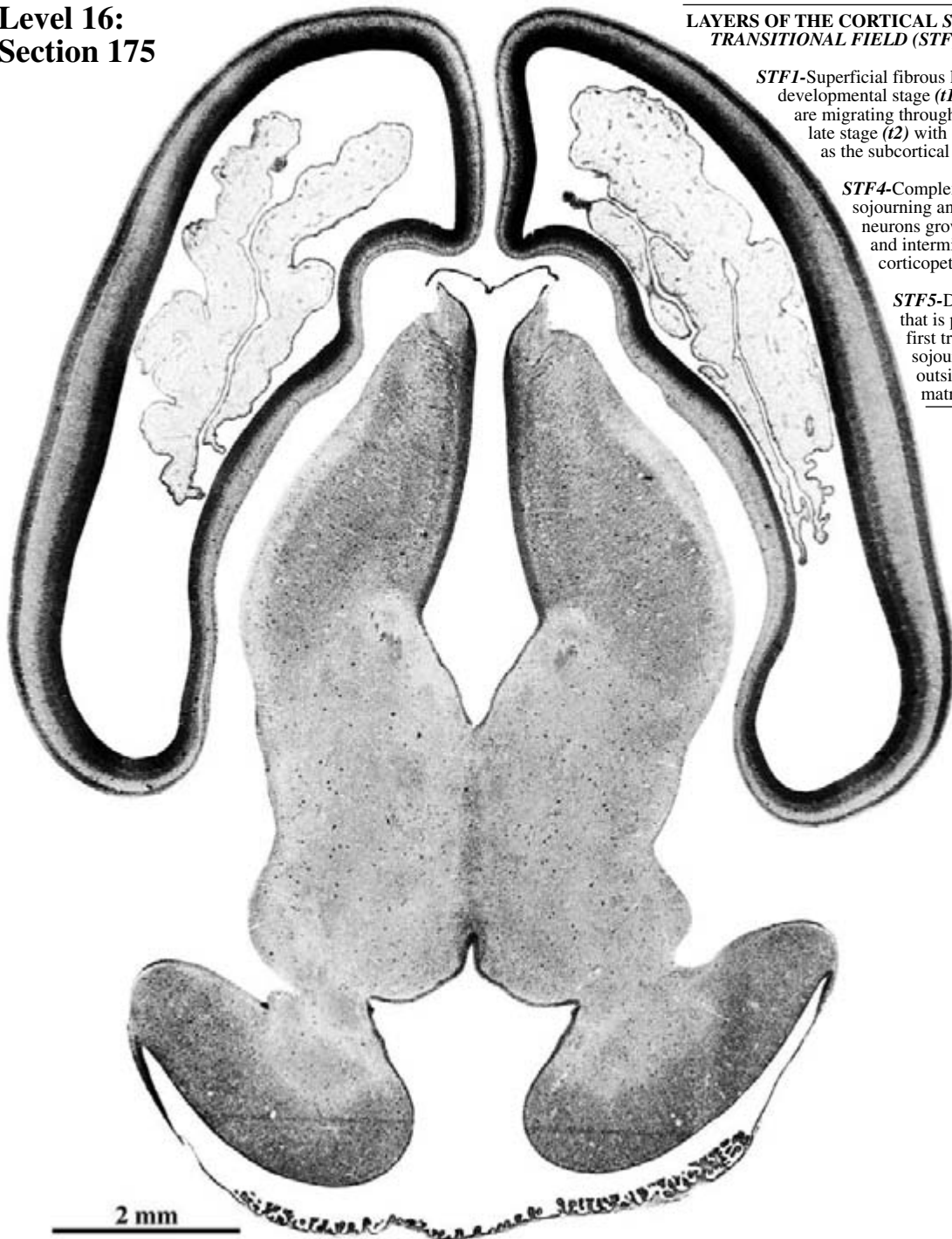
GW9 Coronal

CR 42 mm

M841

Level 16:

Section 175



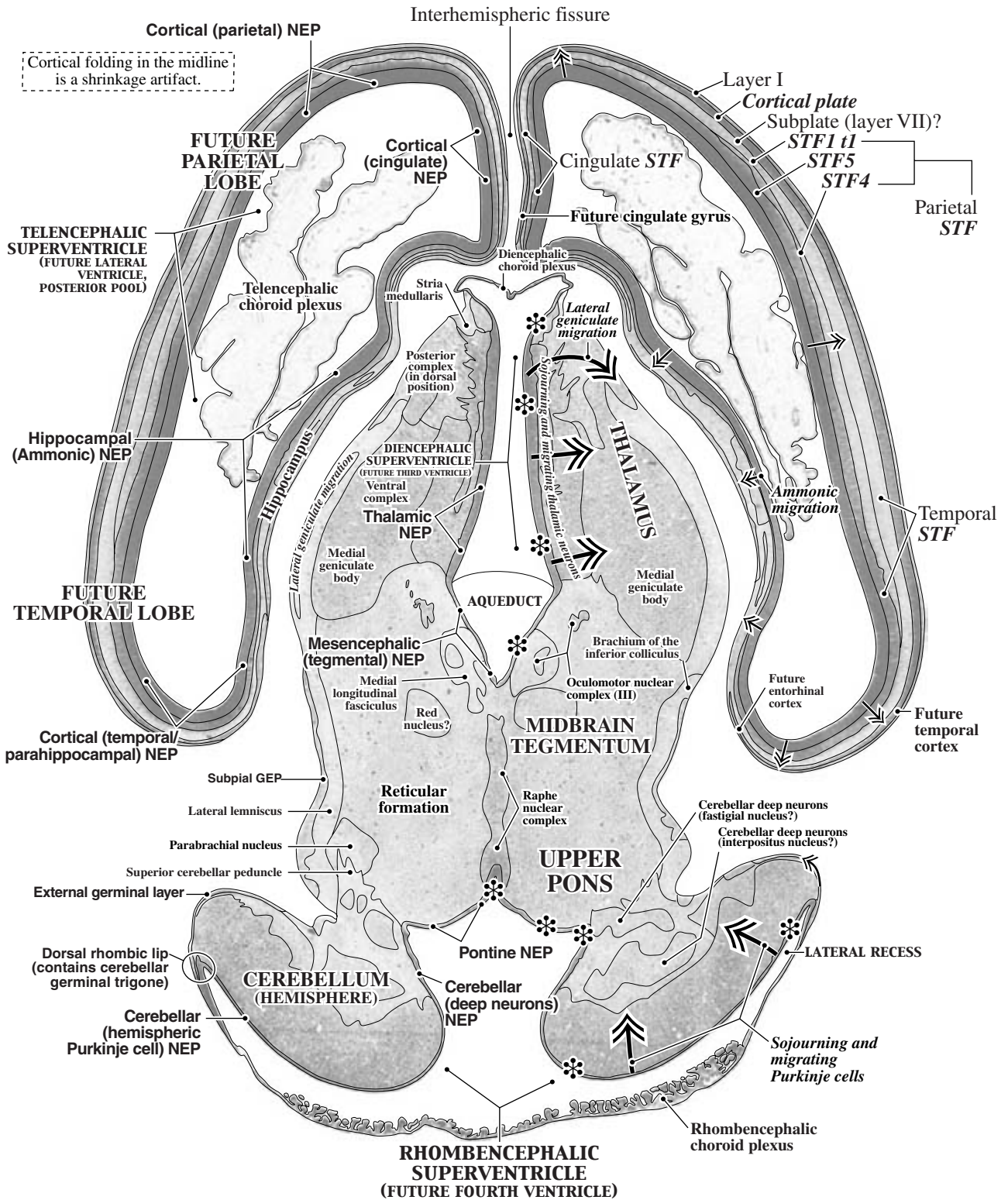
**LAYERS OF THE CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1-Superficial fibrous layer with an early developmental stage (**1**) when many cells are migrating through it, followed by a late stage (**2**) with sparse cells. Endures as the subcortical white matter.

STF4-Complex middle layer where sojourning and migrating cortical neurons grow corticofugal axons and intermingle with corticopetal axons.

STF5-Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 116A

GW9 Coronal

CR 42 mm

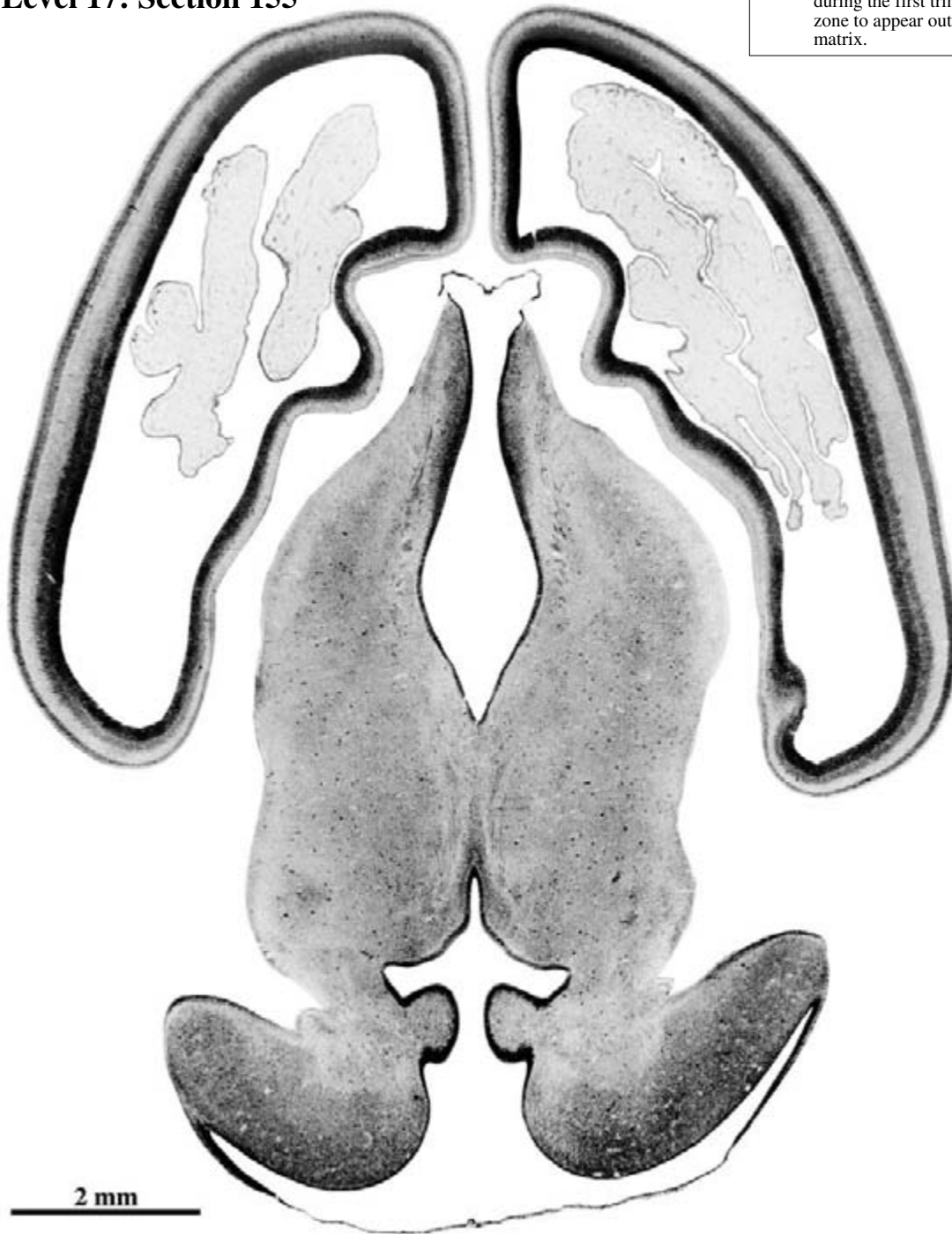
M841

Level 17: Section 155

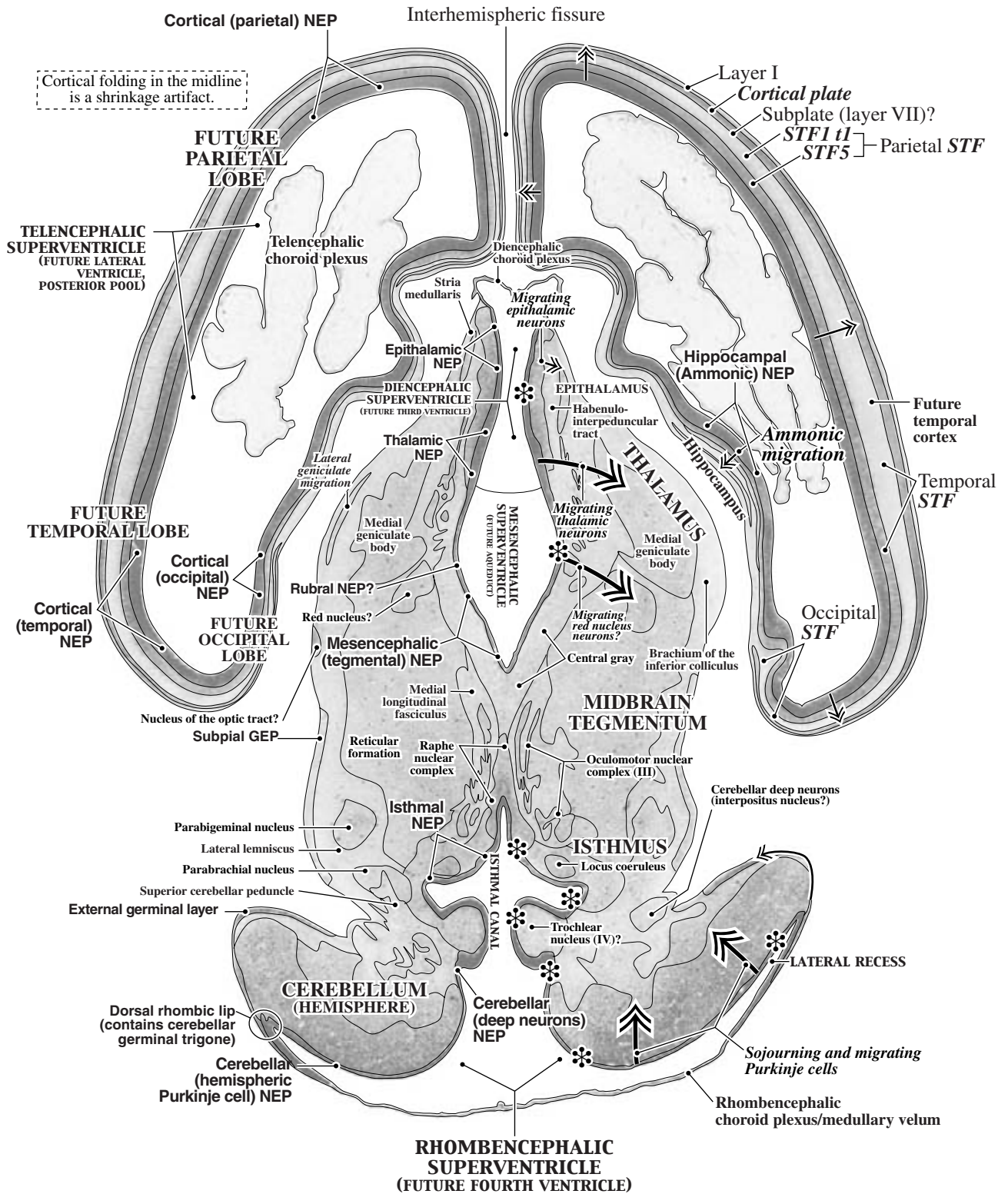
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*I1*) when many cells are migrating through it, followed by a late stage (*I2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioeptithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 117A

GW9 Coronal

CR 42 mm

M841

Level 18: Section 135

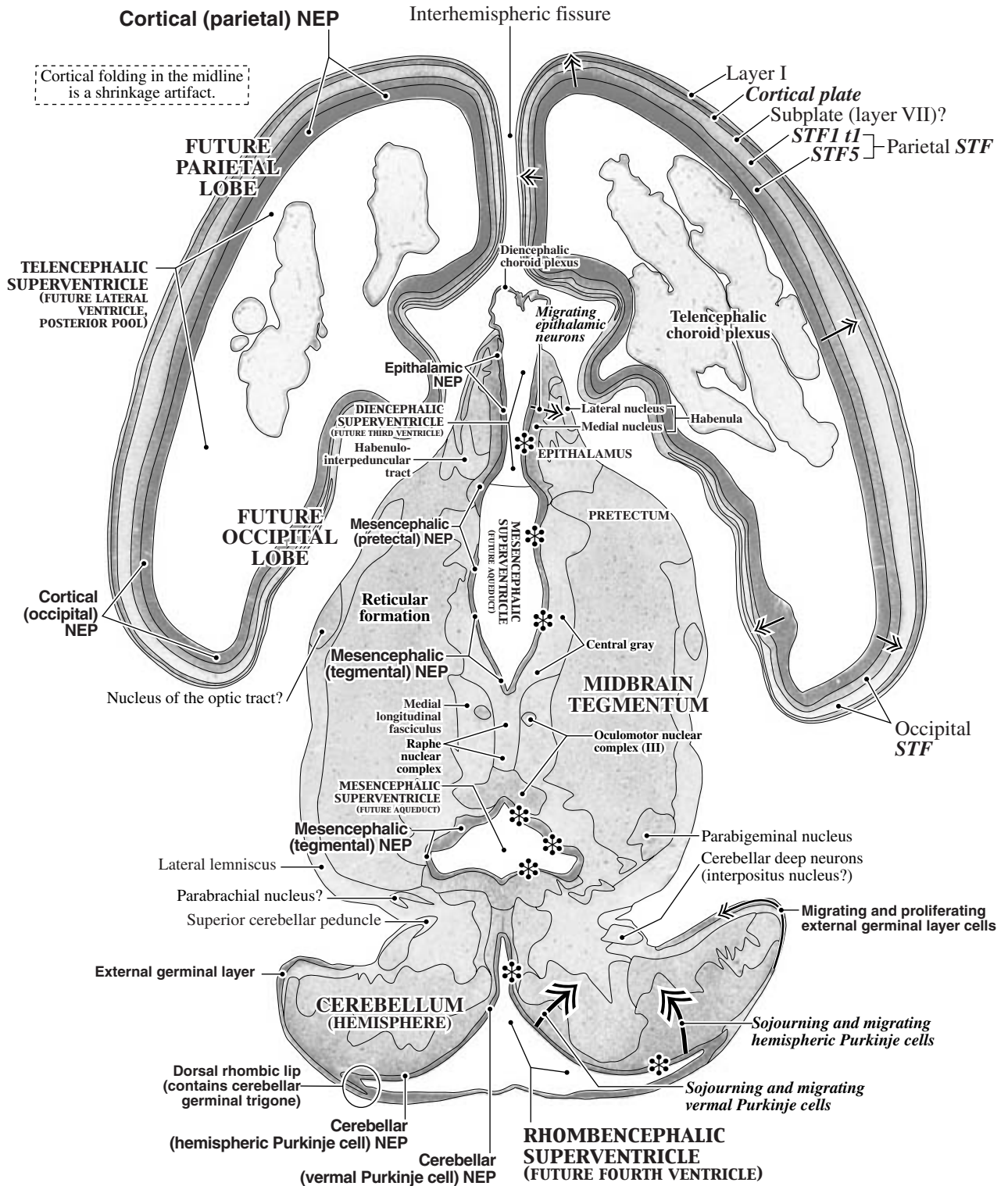
LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



Cortical folding in the midline is a shrinkage artifact.

Layer I
Cortical plate
Subplate (layer VII?)
STF1 t1
STF5 } Parietal STF

Occipital STF

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

NEP - Neuroepithelium

PLATE 118A

GW9 Coronal

CR 42 mm

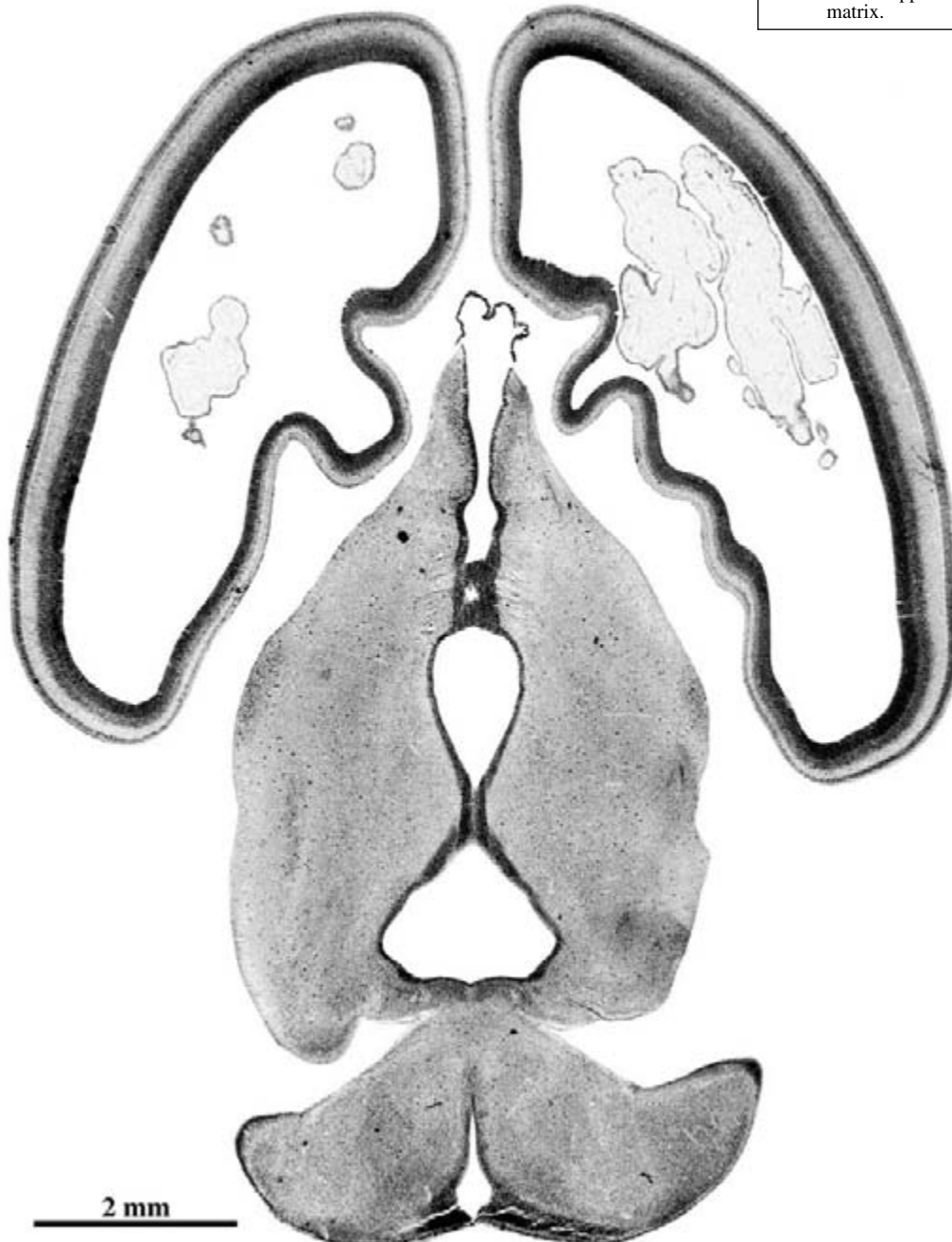
M841

Level 19: Section 124

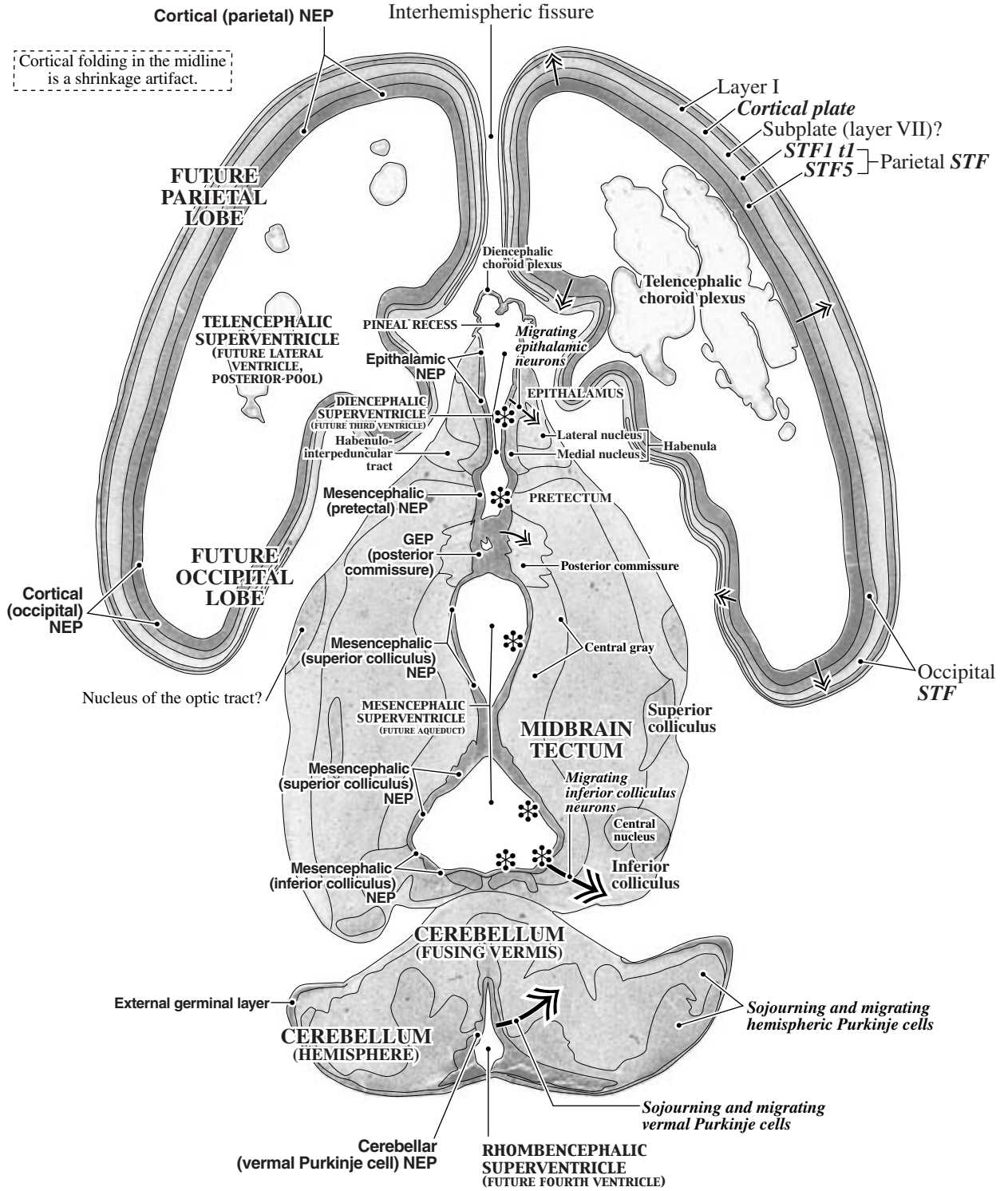
LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



Cortical folding in the midline is a shrinkage artifact.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 119A

GW9 Coronal

CR 42 mm

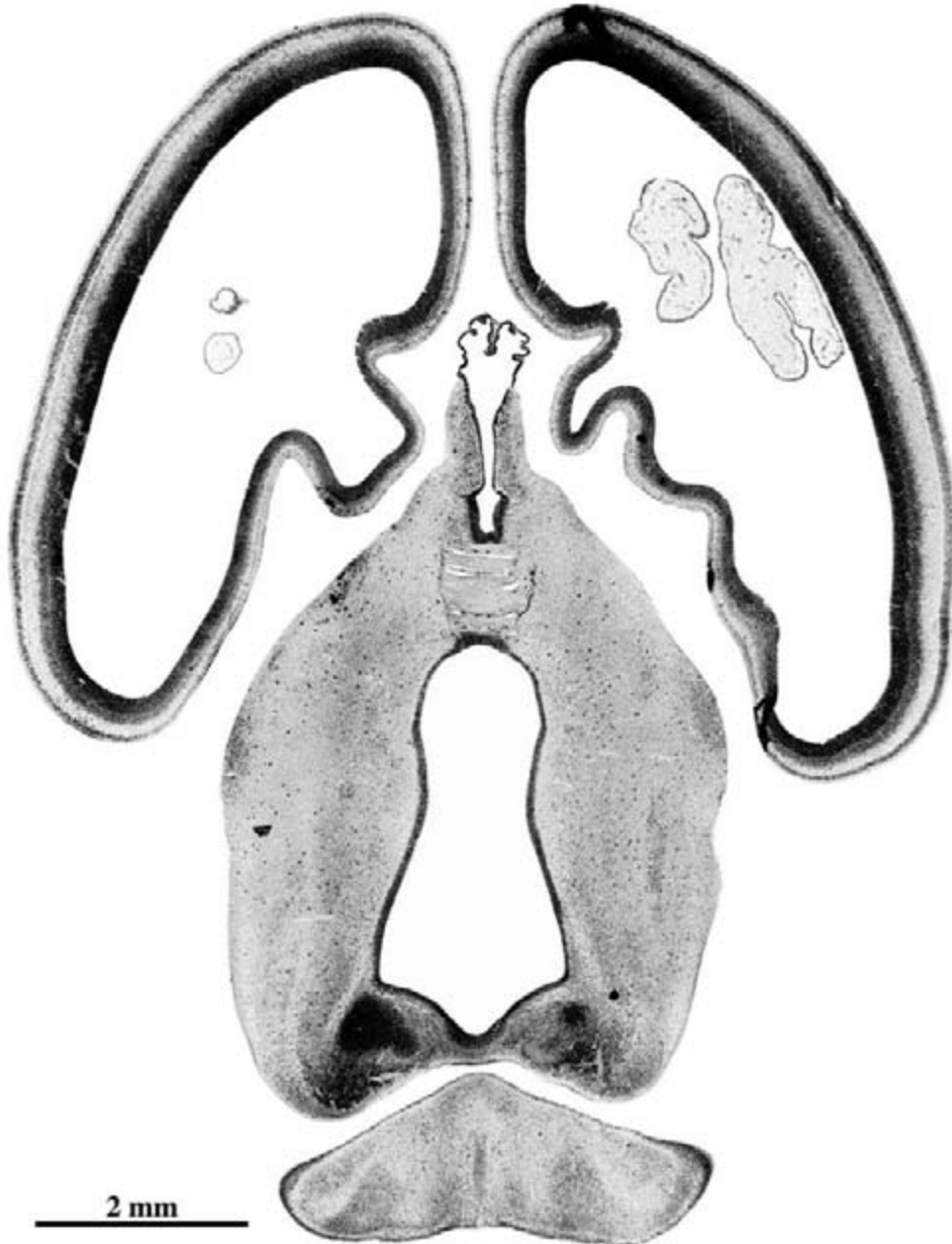
M841

Level 20: Section 112

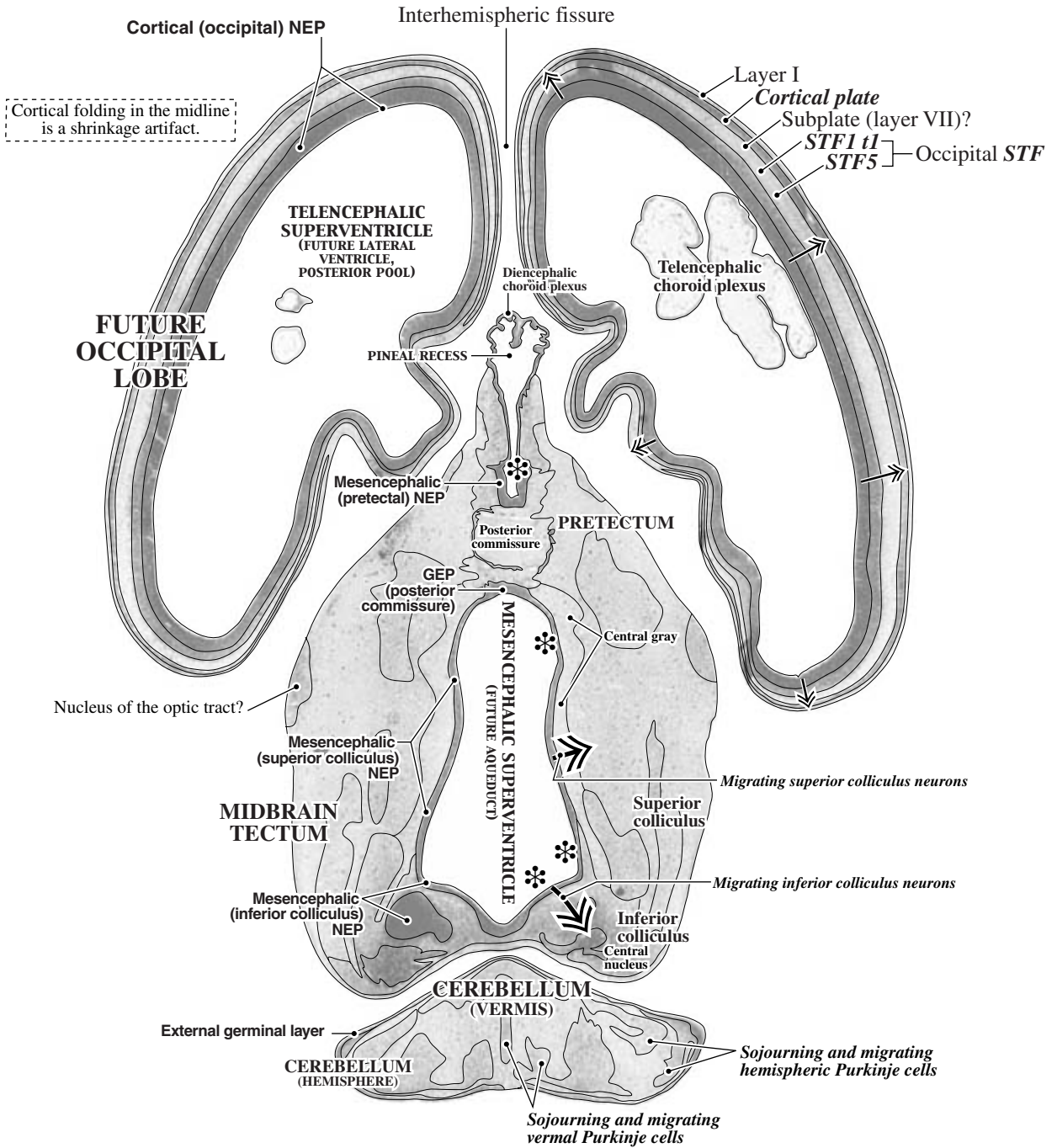
LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



Cortical folding in the midline is a shrinkage artifact.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
 GEP - Glioeptithelium
 NEP - Neuroepithelium

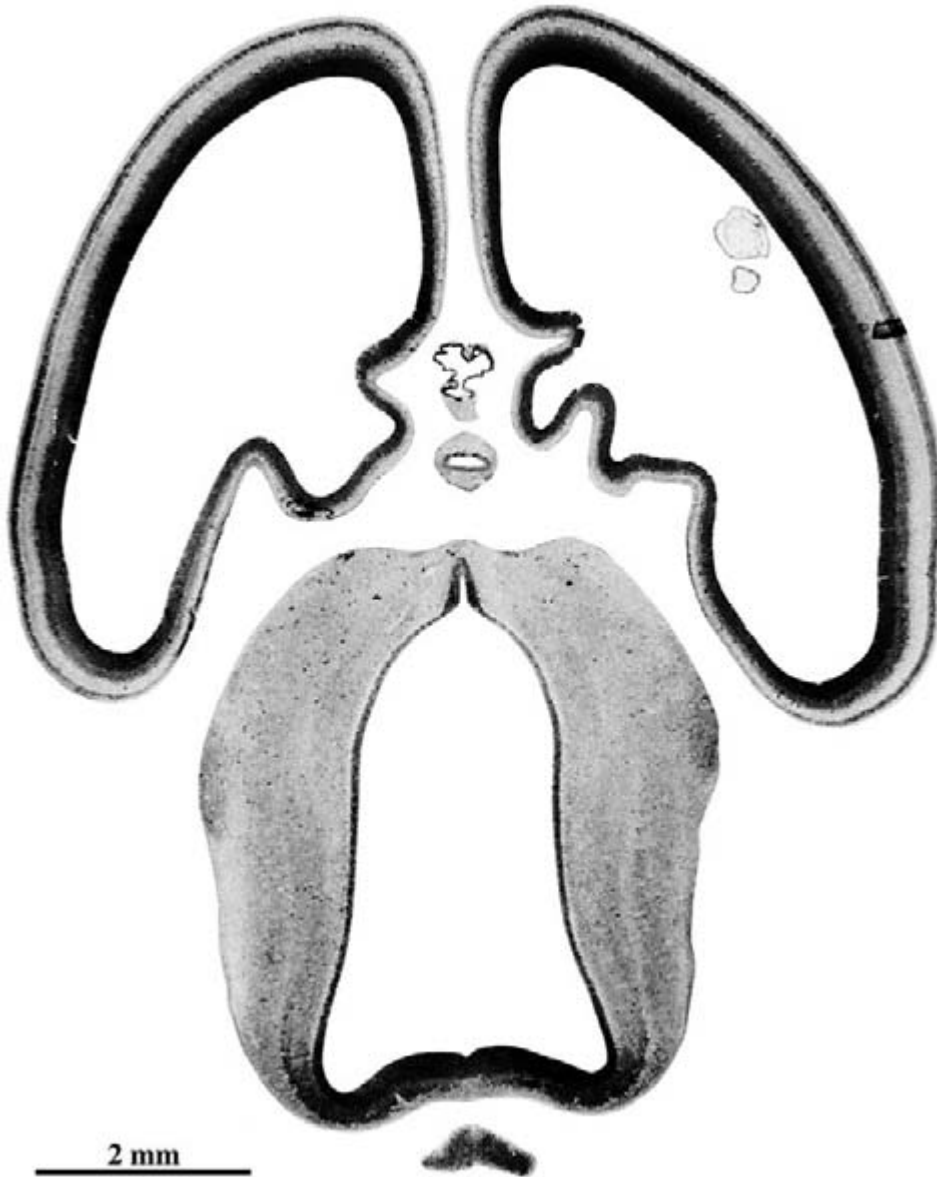
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 120A

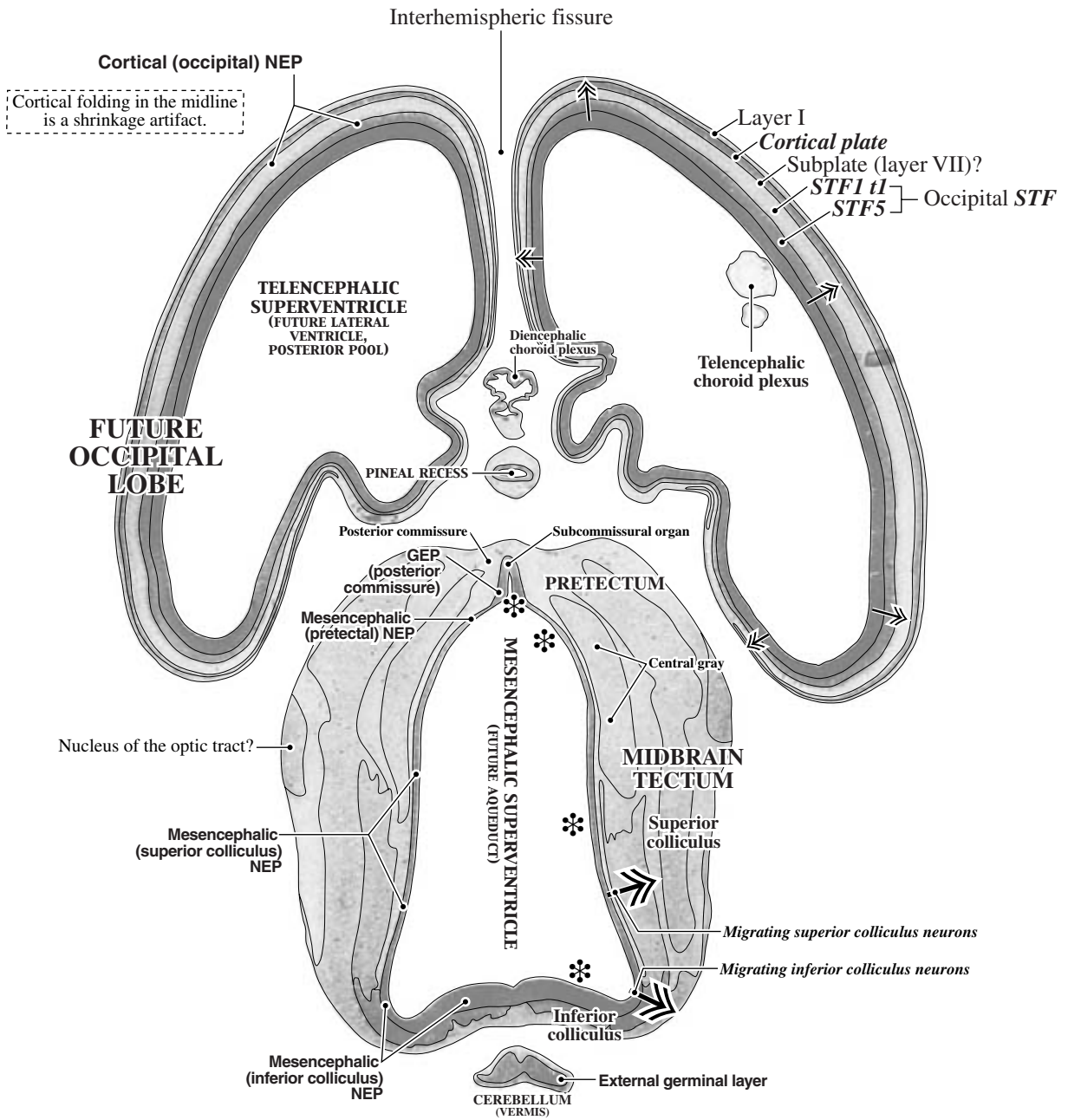
GW9 Coronal
CR 42 mm
M841
Level 21: Section 96

LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



Cortical folding in the midline is a shrinkage artifact.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 121A

GW9 Coronal

CR 42 mm

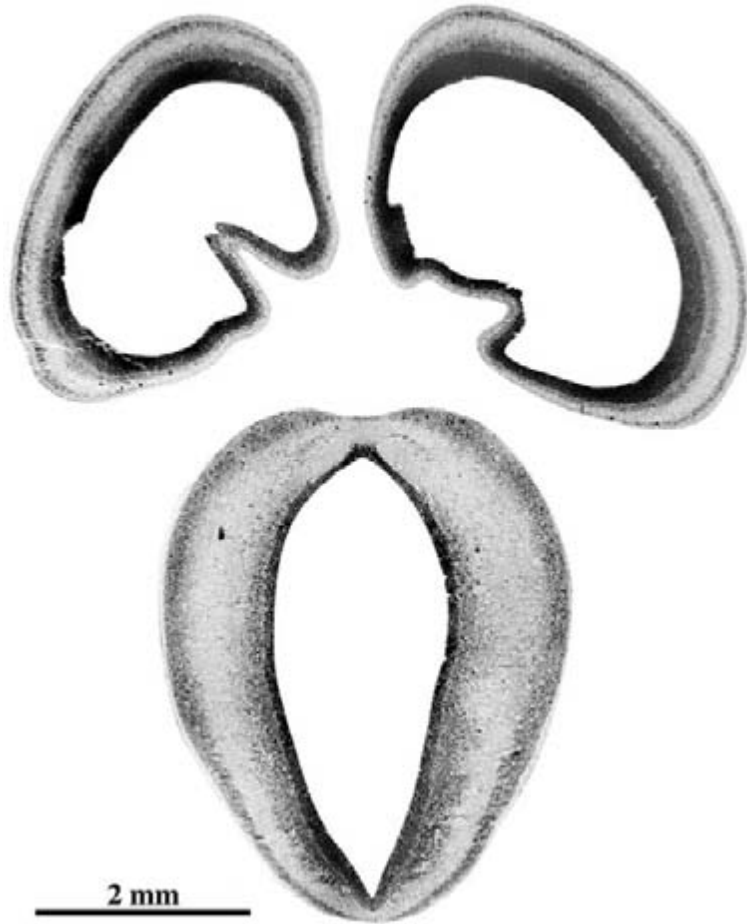
M841

Level 22: Section 39

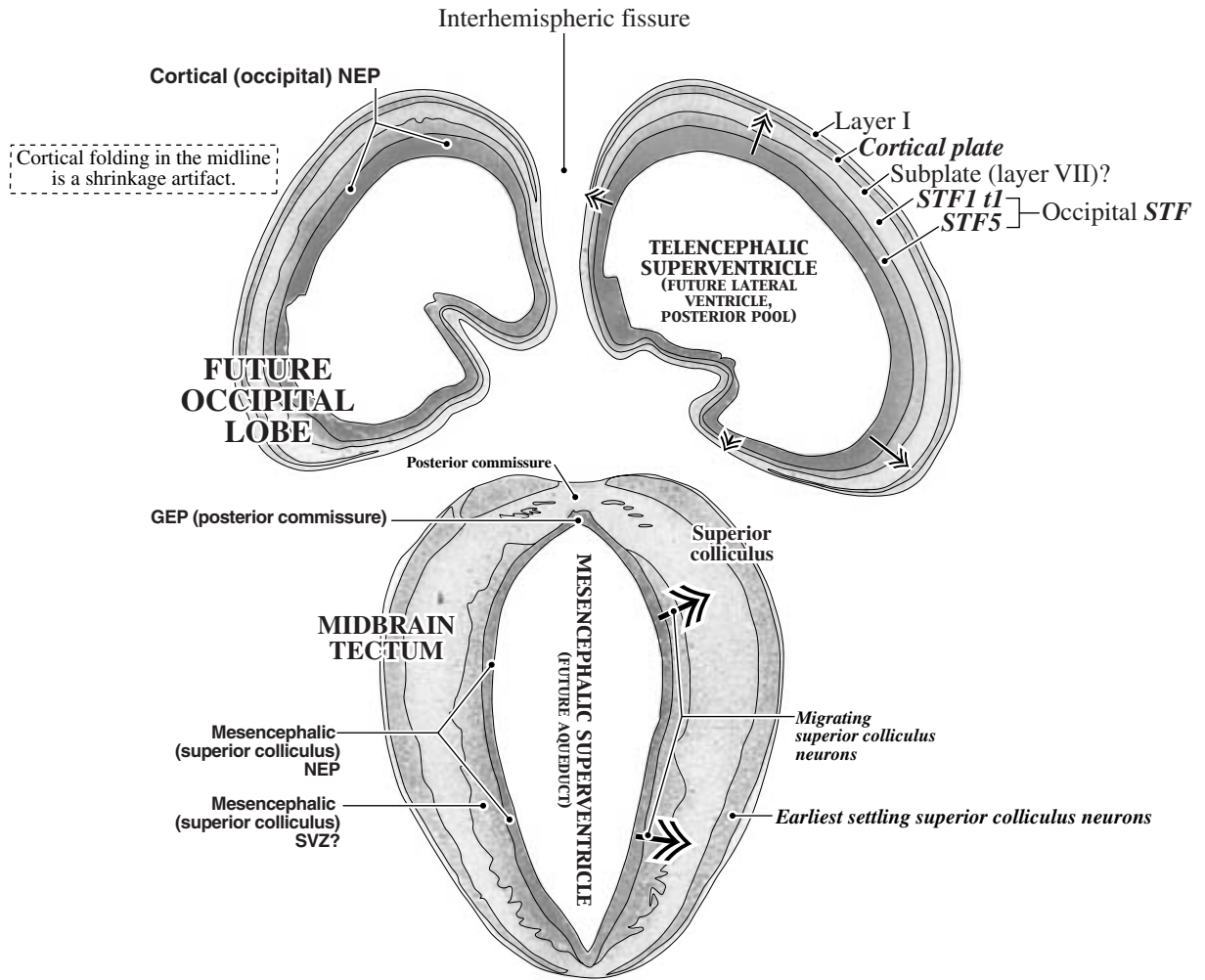
LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (***1***) when many cells are migrating through it, followed by a late stage (***2***) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PART VII: GW8 SAGITTAL

This is specimen number 145 in the Carnegie Collection, designated here as C145, a male with a crown-rump length (CR) of 33 mm estimated to be at gestational week (GW) 8. The entire fetus was cut in the sagittal plane. Information on the date of specimen collection, fixative, section thickness, and embedding medium was not available to us. The sections are thick (probably between 50 to 100 μm) and appear to be embedded in celloidin. Since there is no photograph of C145's brain before histological processing, a specimen from Hochstetter (1919) that is comparable in age to C145 is used to show external brain features at GW8 (**A, Figure 6**). C145's brain structures are easier to understand because sections are closely parallel to the midline; **Figure 6** shows the approximate slight rotations in horizontal (**B**) and vertical (**C**) dimensions. Photographs of 10 sections (**Levels 1-10**) are illustrated at low magnification in four parts (**Plates 122A-D** through **131A-D**). The **A/B** parts show the brain in place in the skull; the **C/D** parts show only the brain (and some peripheral ganglia) at slightly higher magnification. **Plates 132-147** show high magnification views of various parts of the brain. All of the high-magnification plates are rotated 90° (landscape orientation) to show photographs at higher magnification in the available page space.

C145 is considerably less mature than the GW9 specimens. One of the most notable features of this specimen is the larger volume of the brain ventricles when compared to the brain parenchyma (areas where neurons migrate, settle, and differentiate). The largest structure in each of the brain's major subdivisions is the *superventricles* in their cores. For example, the telencephalon is largely occupied by the telencephalic superventricle. The thickness of the parenchyma is a key to the degree of maturation of the various brain structures. It is thickest in the medulla, pons, and midbrain tegmentum where most of the neurons have been generated and thinner in the cerebellum, midbrain tectum, and diencephalon. Within the telencephalon, the cerebral cortex has a very thin parenchyma, while the basal telen-

cephalon and parts of the basal ganglia have thick parenchymal components.

Throughout the cerebral cortex, the *neuroepithelium* is prominent as the sole germinal matrix. The *stratified transitional field (STF)* contains *STF1* and *STF5* only in lateral areas. The pronounced anterolateral (thicker) to dorsomedial (thinner) maturation gradient is evident in both the *cortical plate* and the *STF* layers. The olfactory bulb is just beginning to evaginate in front of the basal telencephalic neuroepithelium. Neurons are just beginning to migrate in the hippocampus. A massive *neuroepithelium/subventricular zone* overlies the amygdala, nucleus accumbens, and striatum (caudate and putamen) where neurons (and glia) are being generated.

The cerebellum has a thicker *neuroepithelium*, indicating many Purkinje cells are being generated. Earlier-generated Purkinje cells are sojourning in a dense layer outside the neuroepithelium. Most of the deep neurons are superficial in the cerebellum. The *external germinal layer (egl)* is barely visible emanating from the germinal trigone in the dorsal rhombic lip.

In sections near the midline, the brainstem *neuroepithelium* varies in thickness. It is thinner in the midbrain tegmentum, pons, and medulla in accordance with an earlier maturation gradient. Most neurons have been generated in these structures and are settling. In the cerebellum (see above), midbrain tectum, and diencephalon, the neuroepithelium is thicker indicating that substantial neurogenesis is happening in these structures, but many neurons have already been produced. However, more lateral sections have a thick *precerebellar neuroepithelium* in the medulla. Since the pontine gray is totally absent, nearly all of those neurons have yet to be generated in the precerebellar neuroepithelium; some of the earliest-generated pontine gray neurons are migrating in the *anterior extramural migratory stream*.

GW8 SAGITTAL

A perfect sagittal cut through the brain bisects the cerebral cortex into two separate hemispheres by passing through the interhemispheric fissure, and does the same in the brainstem by passing through the midline of the ventricles.

Sections of C145's brain are very close to the midline both horizontally (+3.48°, top view) and vertically (+3.82°, back view). In each section illustrated on the following pages, the anterior edge of the cortex (top right) is tilted toward the observer, while the medulla and upper spinal cord (bottom) are tilted away from the observer.

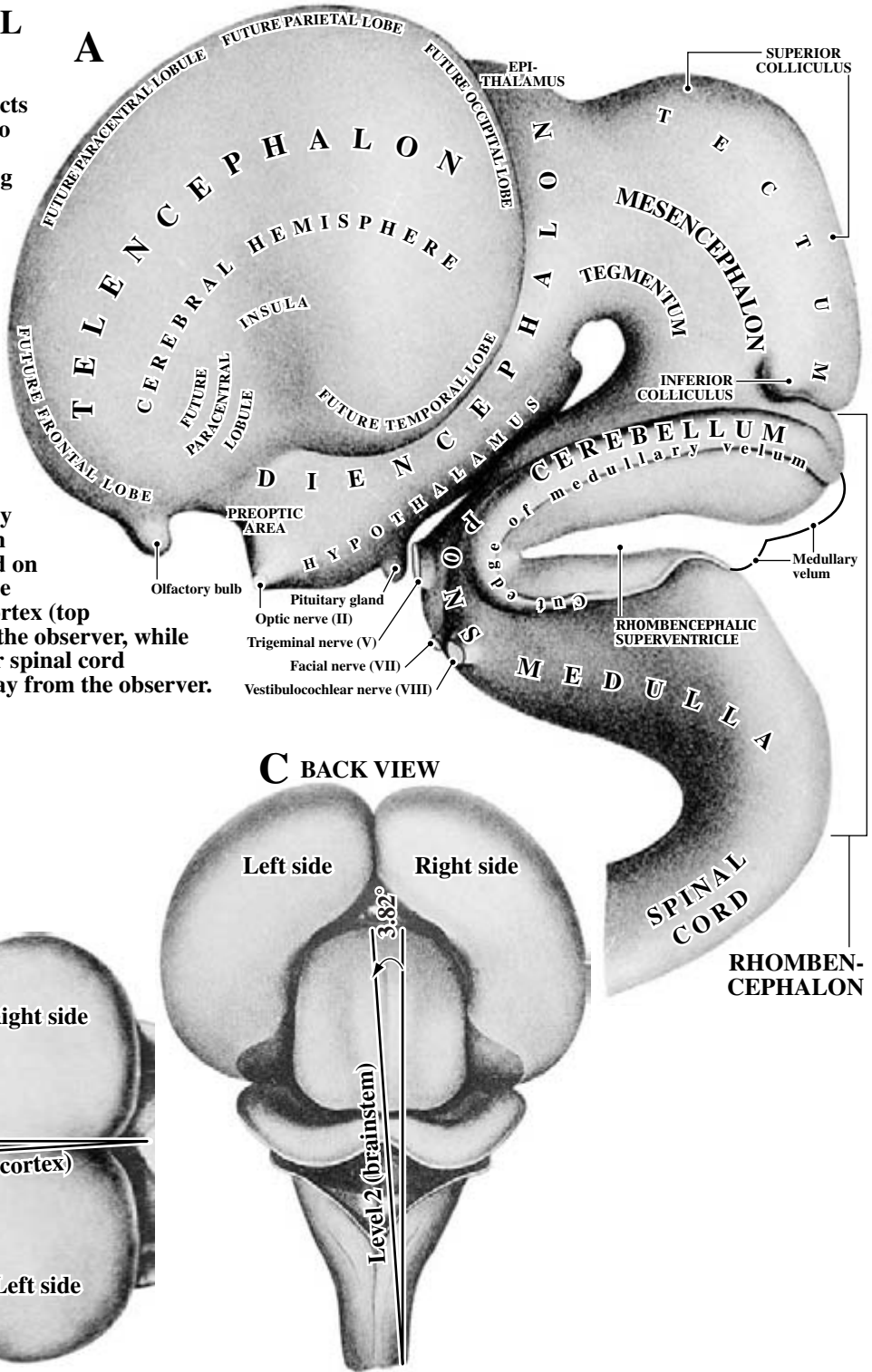
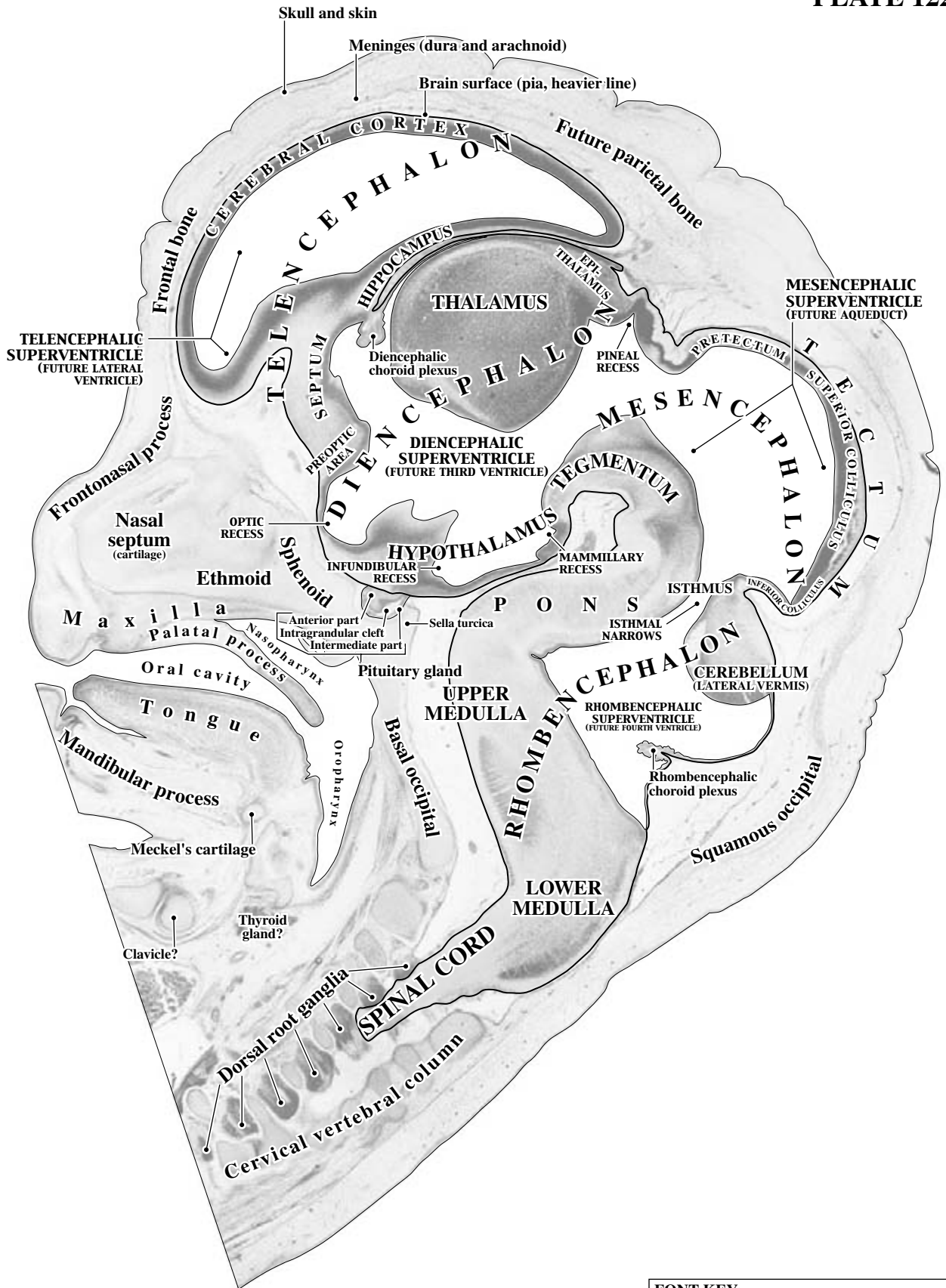


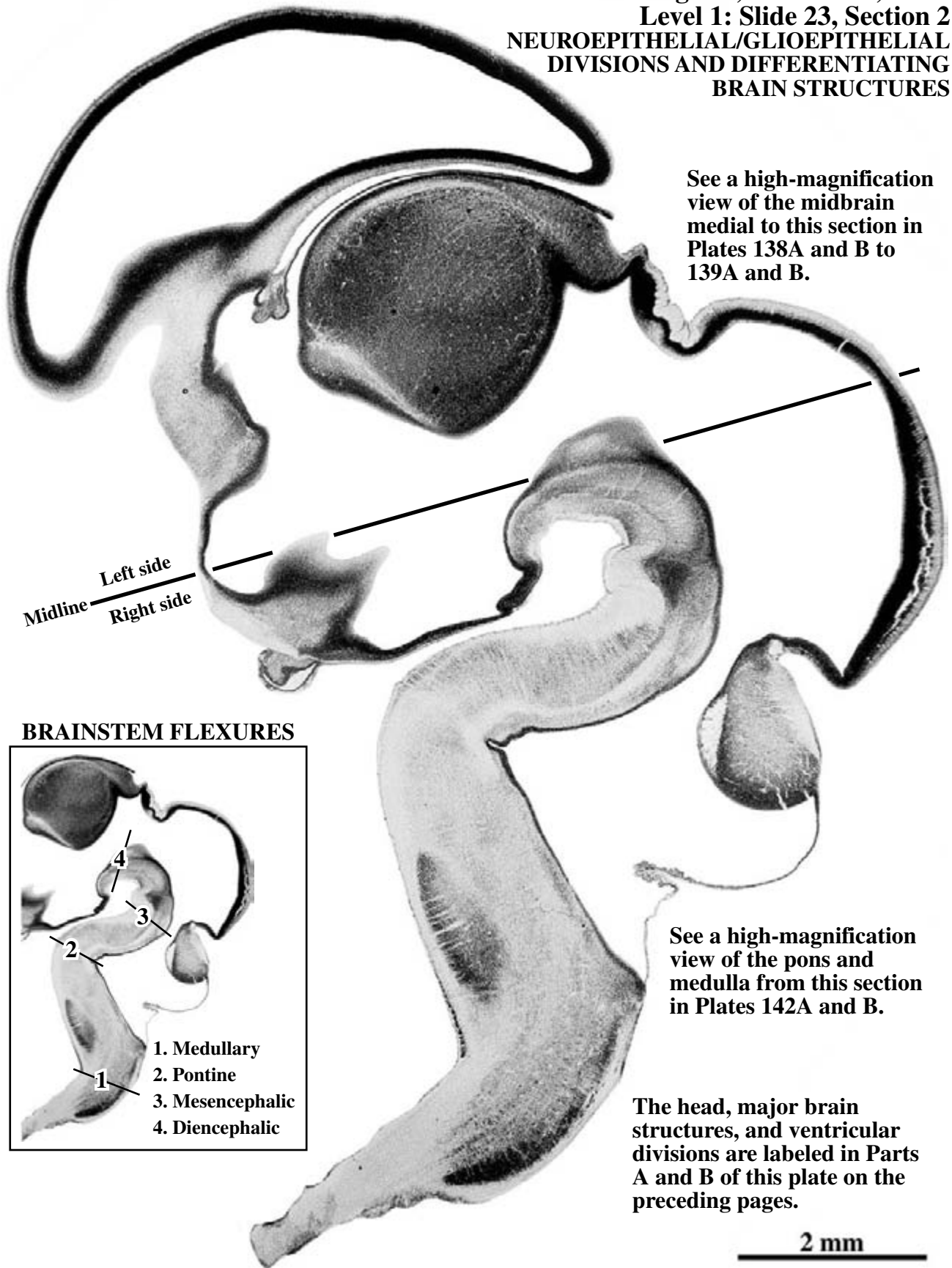
Figure 6. A, The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 27 mm (modified from Figure 39, Table VII, Hochstetter, 1919) identifies external features of a brain similar to C145 (CR 33 mm). B, Top view of the brain with a crown-rump length of 38 mm (modified from Figure 45, Table VIII, Hochstetter, 1919) shows how C145's sections rotate from a line parallel to the horizontal midline in the interhemispheric fissure. C, Back view of the brain in B (modified from Figure 44, Table VIII, Hochstetter, 1919) shows how C145's sections rotate from a line parallel to the vertical midline in the brainstem and upper cervical spinal cord.



Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



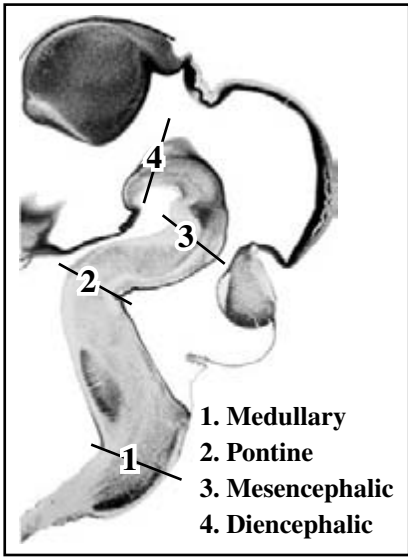
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See a high-magnification view of the midbrain medial to this section in Plates 138A and B to 139A and B.

Left side
Midline
Right side

BRAINSTEM FLEXURES

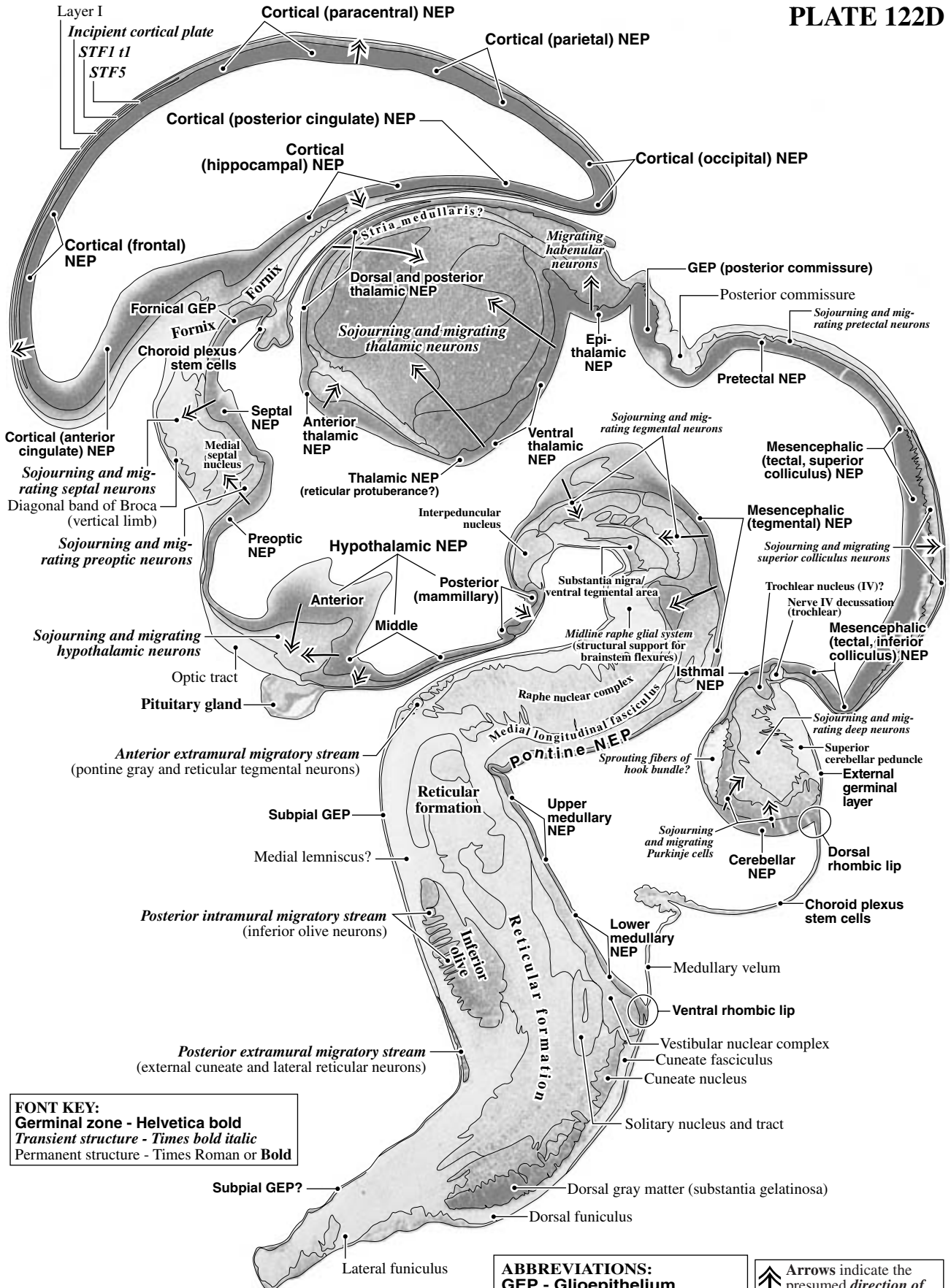


- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

See a high-magnification view of the pons and medulla from this section in Plates 142A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

2 mm



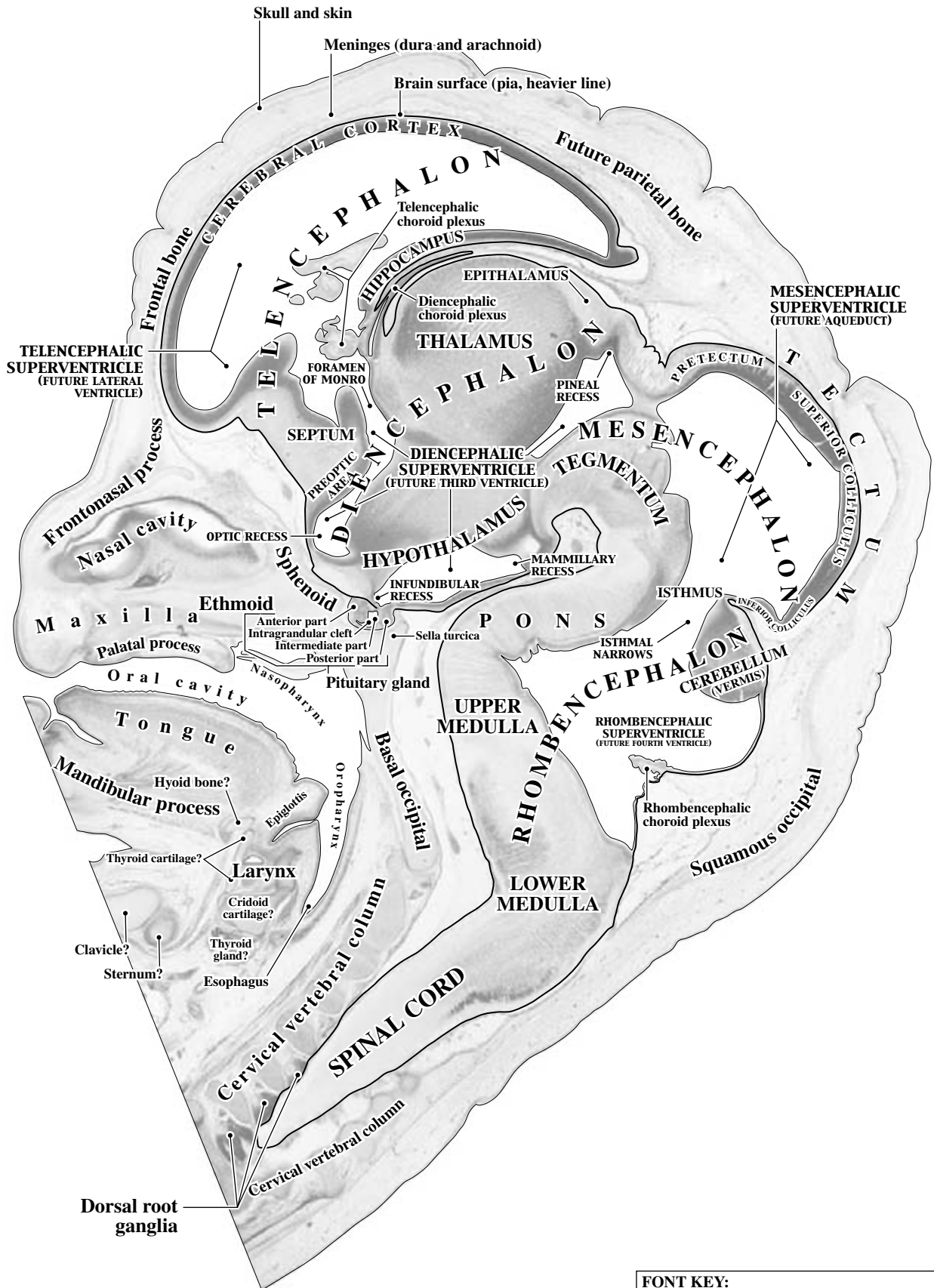
FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
STF - Stratified transitional field

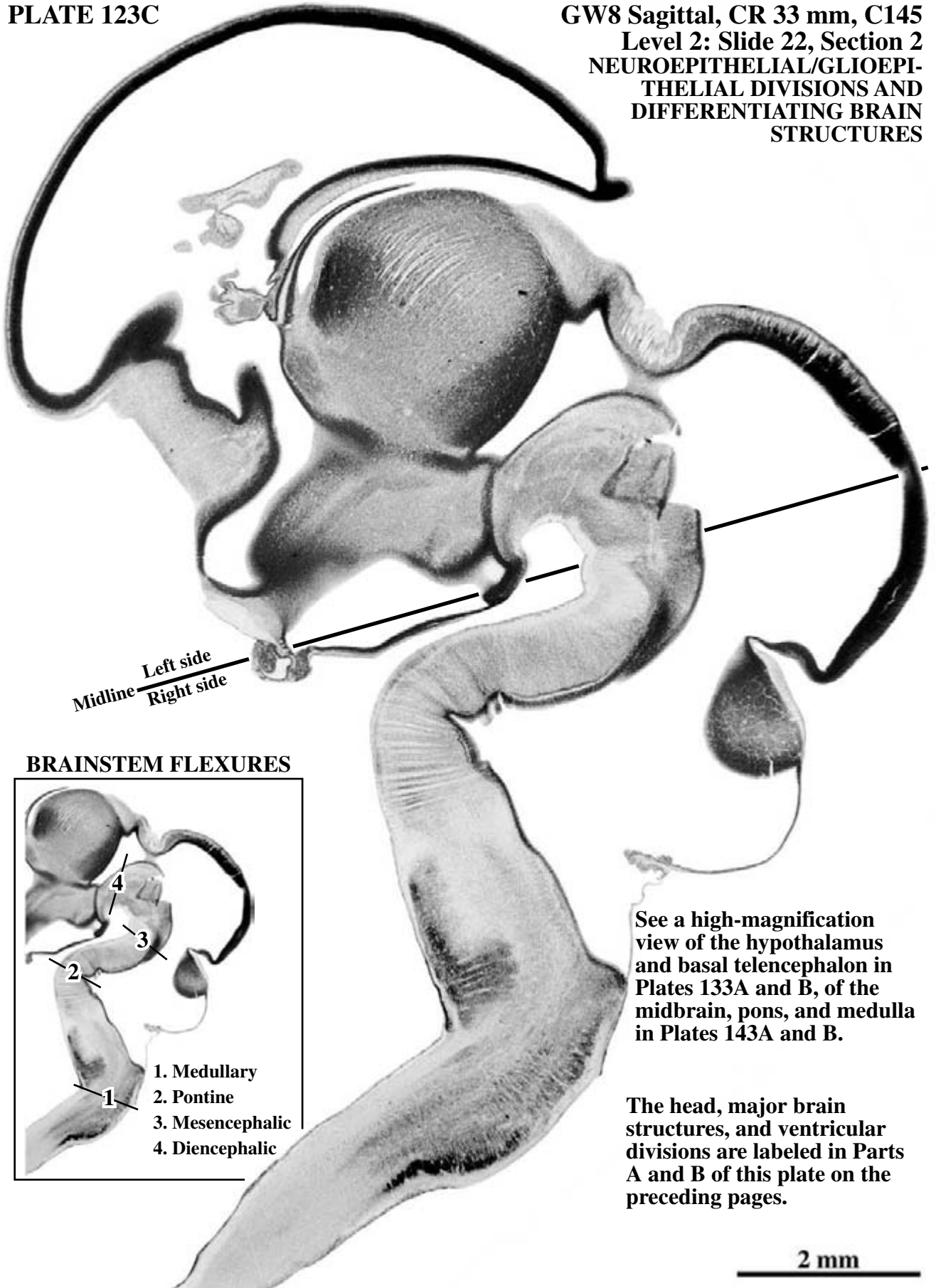
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.



Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

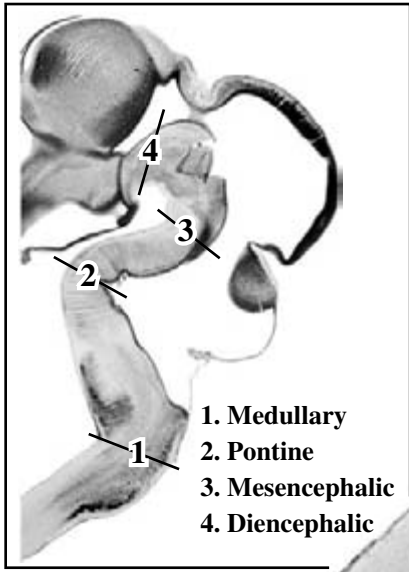


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold



Midline
Left side
Right side

BRAINSTEM FLEXURES

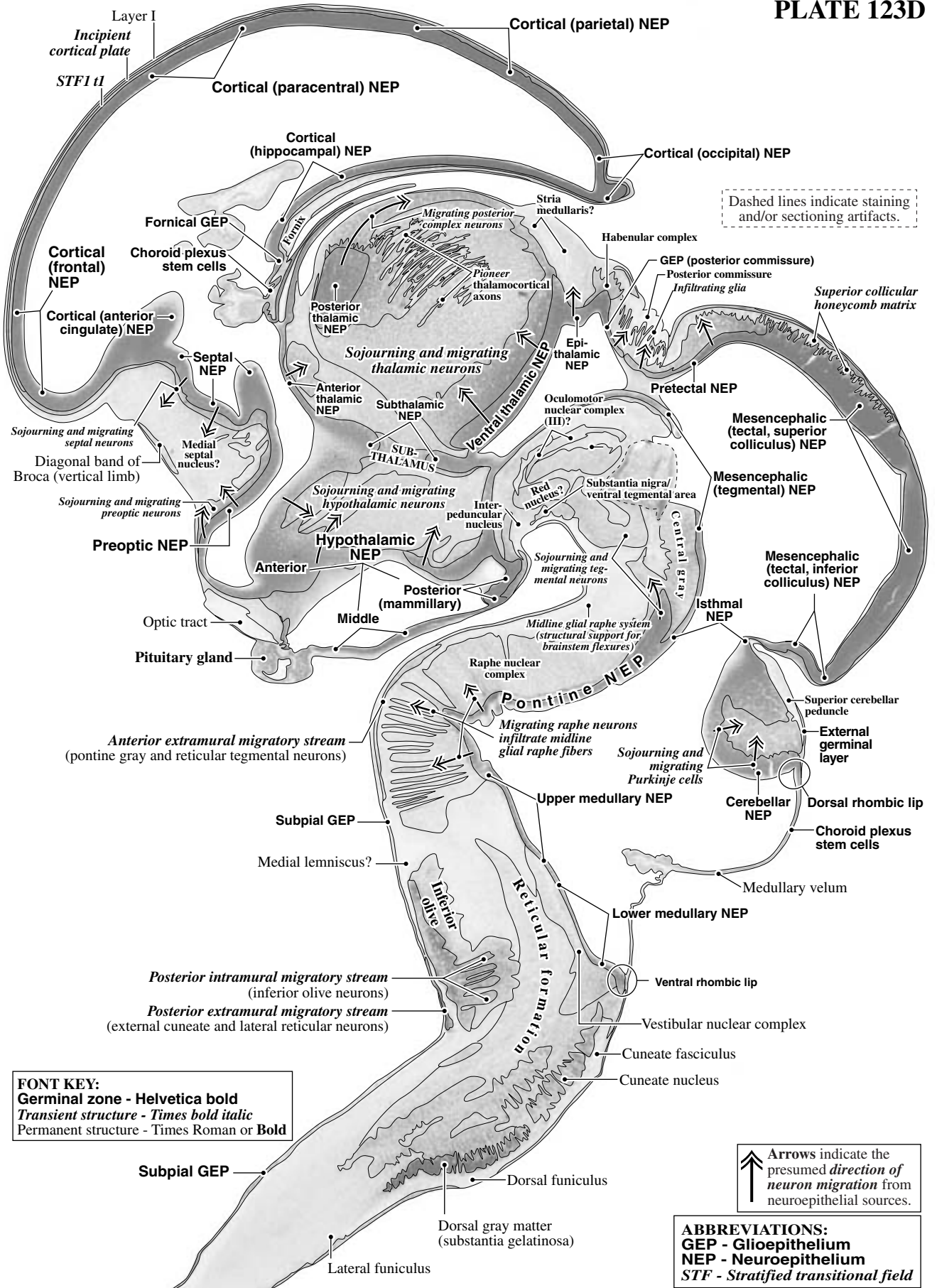


- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

See a high-magnification view of the hypothalamus and basal telencephalon in Plates 133A and B, of the midbrain, pons, and medulla in Plates 143A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

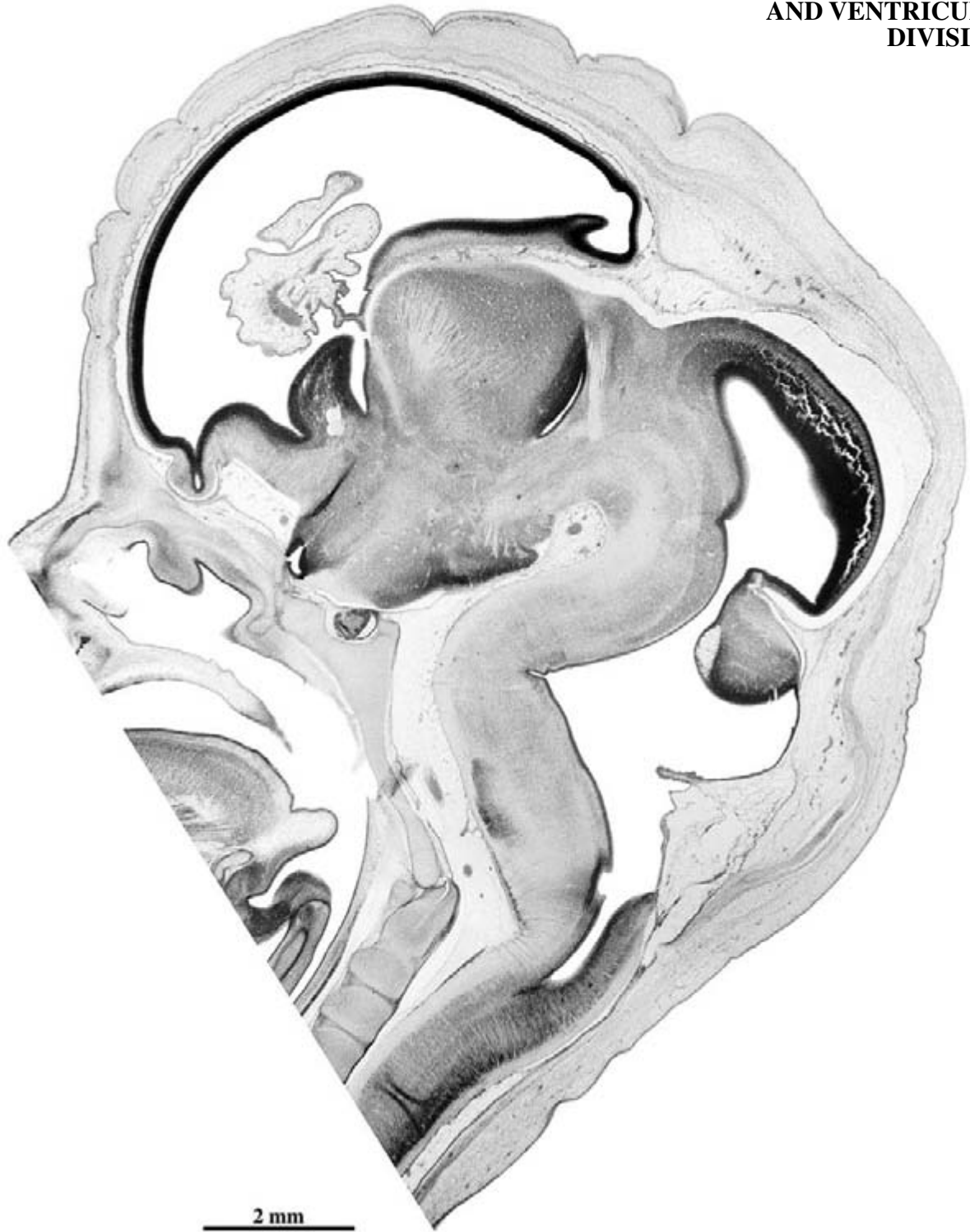
2 mm



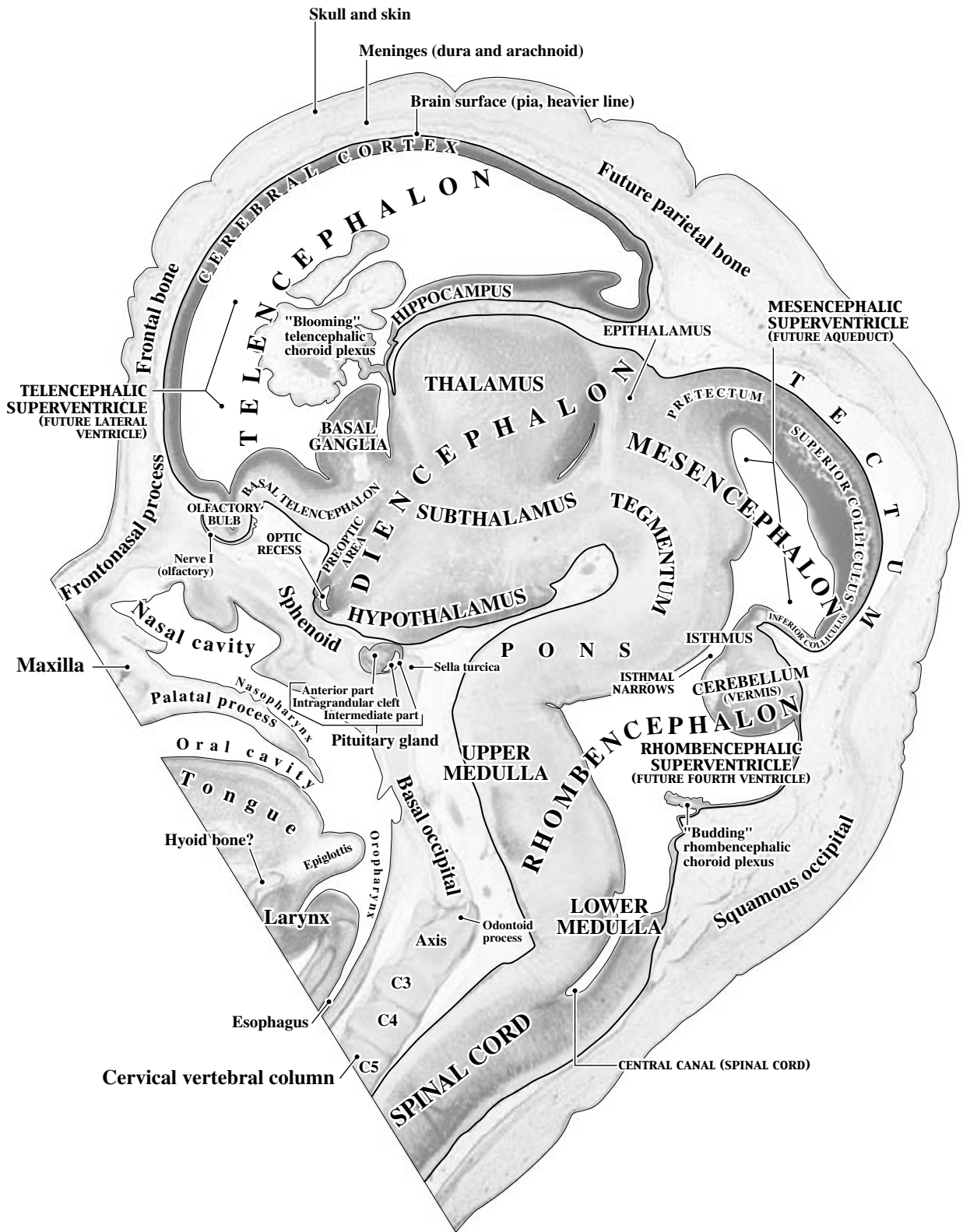
FONT KEY:
 Germinal zone - Helvetica bold
 Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium
 STF - Stratified transitional field



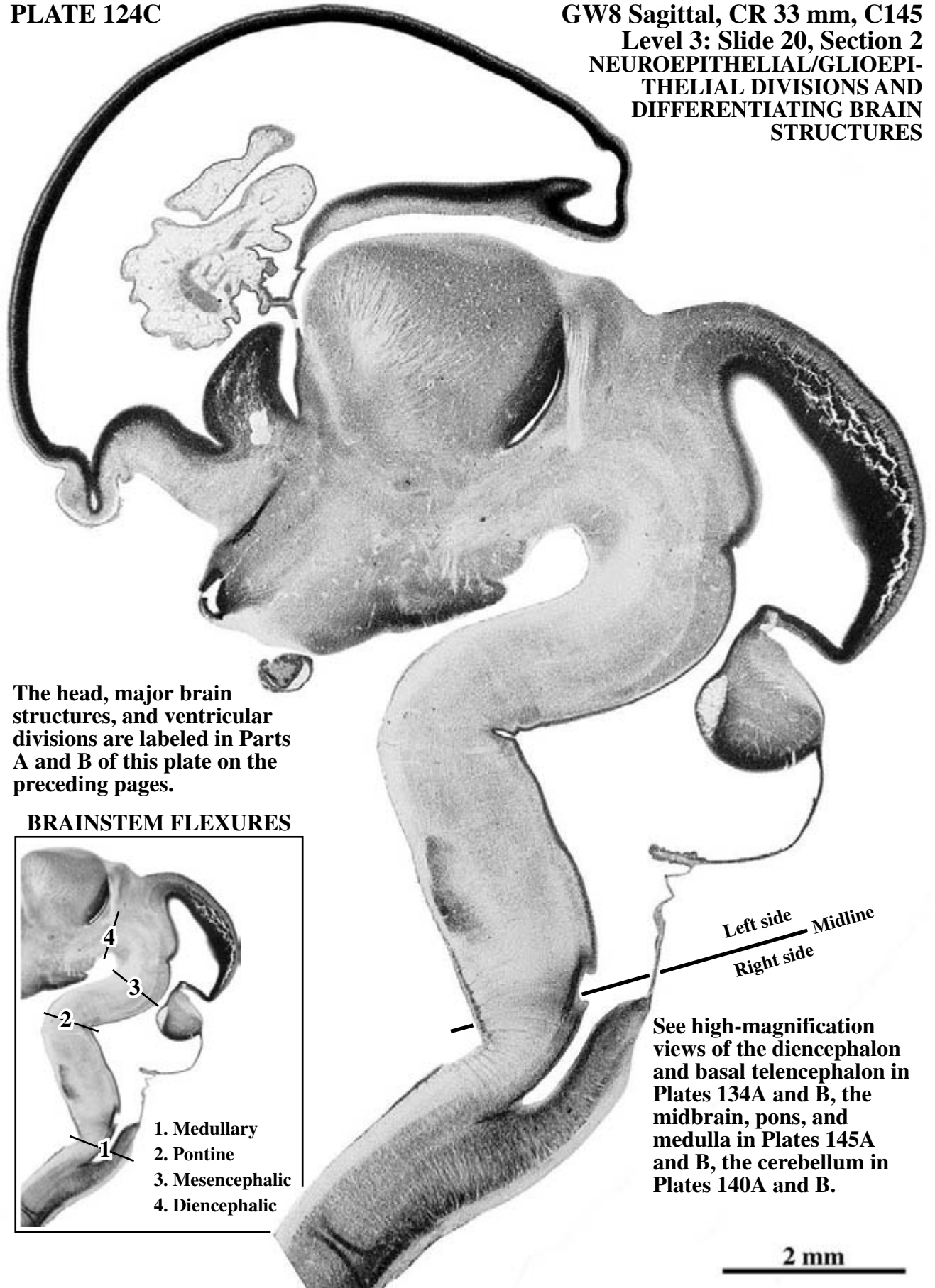
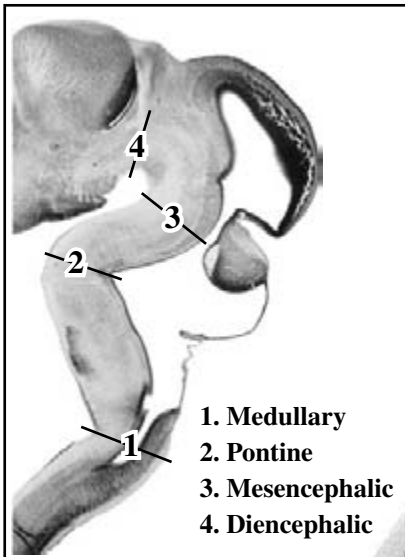
Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

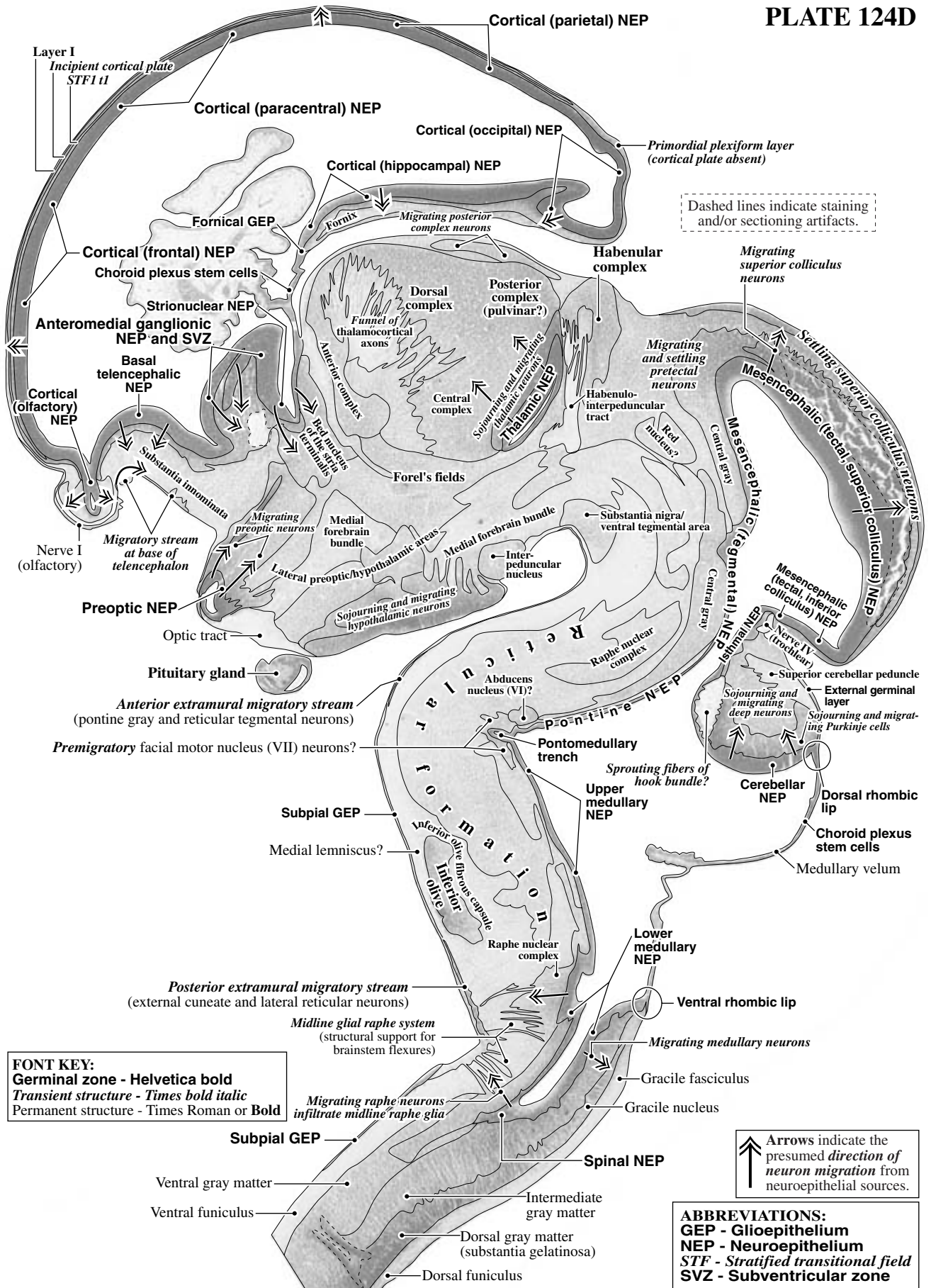
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

BRAINSTEM FLEXURES



See high-magnification views of the diencephalon and basal telencephalon in Plates 134A and B, the midbrain, pons, and medulla in Plates 145A and B, the cerebellum in Plates 140A and B.

2 mm



Dashed lines indicate staining and/or sectioning artifacts.

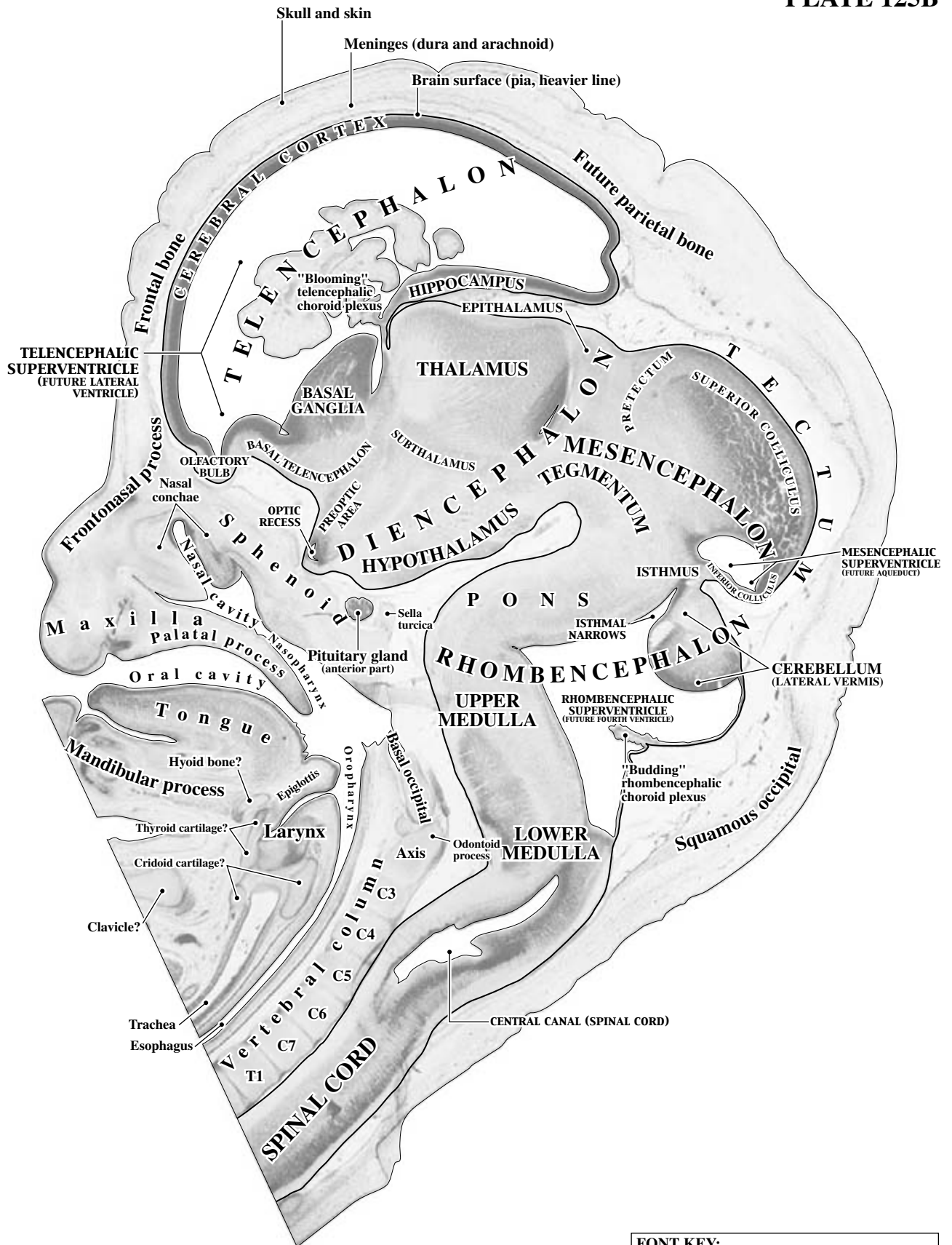
FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

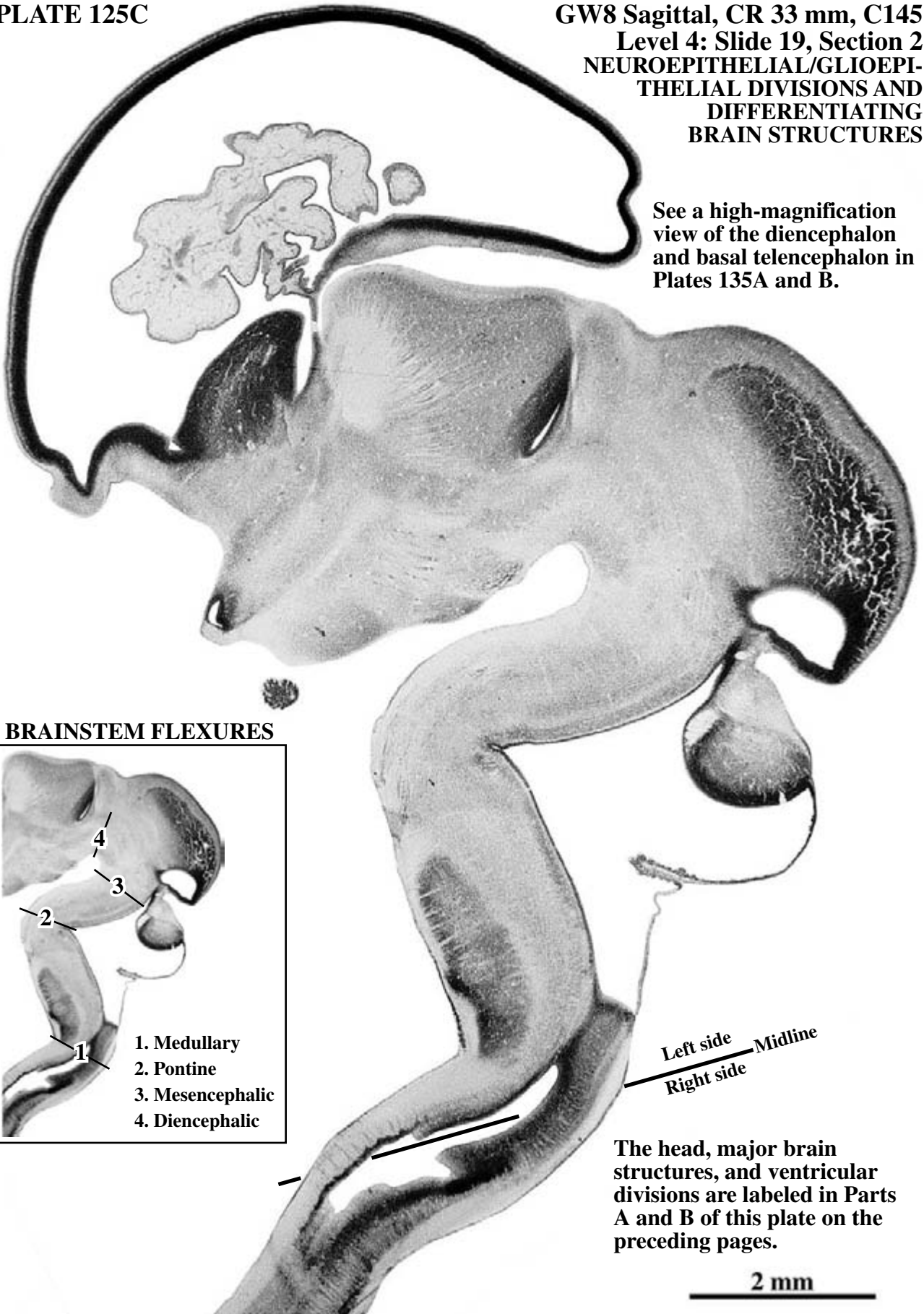


Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

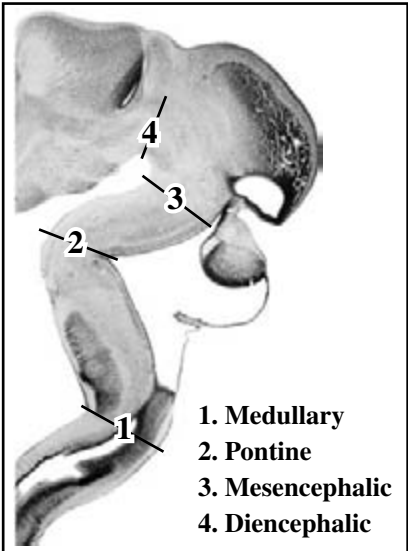


**GW8 Sagittal, CR 33 mm, C145
Level 4: Slide 19, Section 2
NEUROEPITHELIAL/GLIOEPI-
THELIAL DIVISIONS AND
DIFFERENTIATING
BRAIN STRUCTURES**

See a high-magnification
view of the diencephalon
and basal telencephalon in
Plates 135A and B.



BRAINSTEM FLEXURES

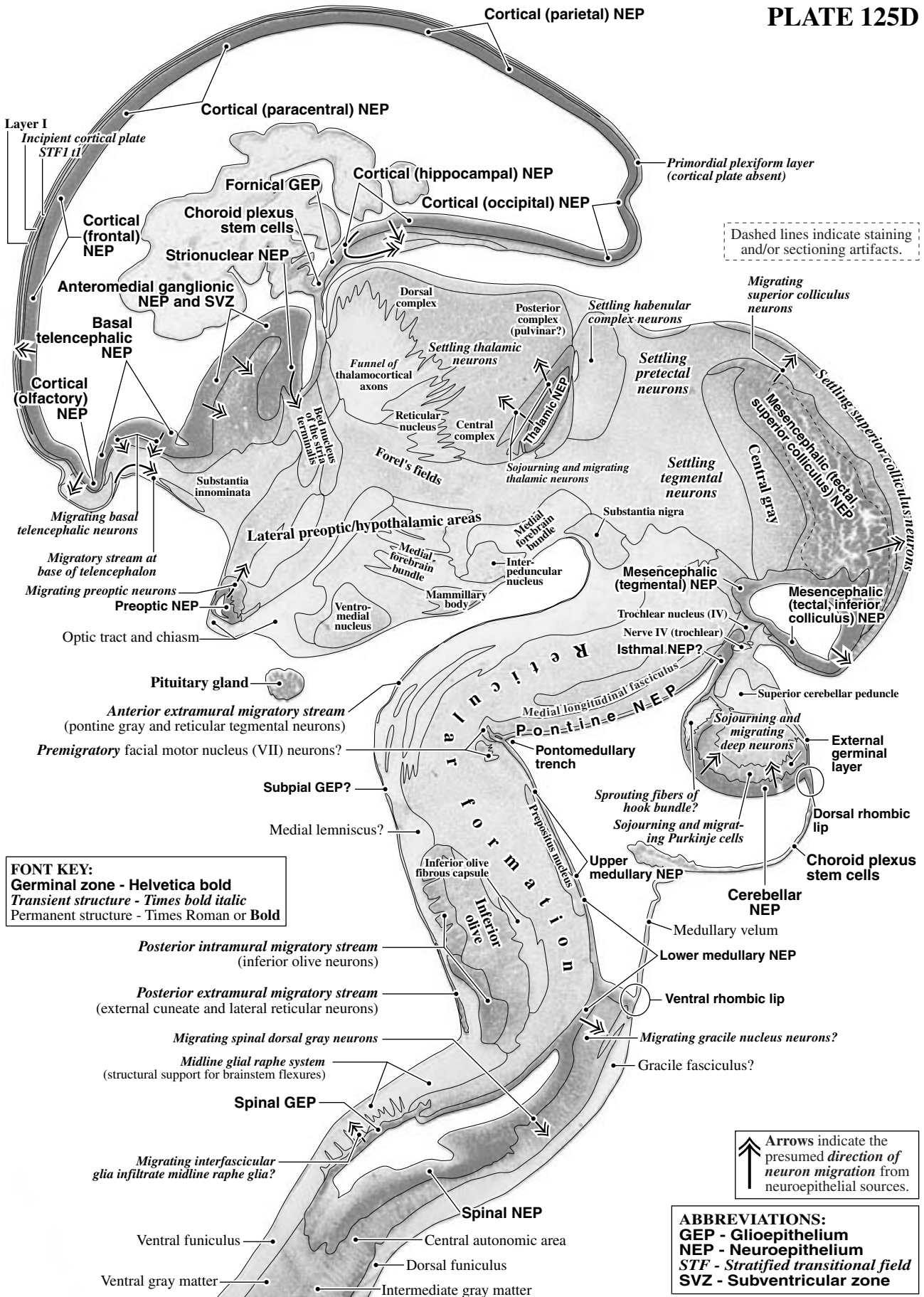


- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

Left side Midline
Right side

The head, major brain
structures, and ventricular
divisions are labeled in Parts
A and B of this plate on the
preceding pages.

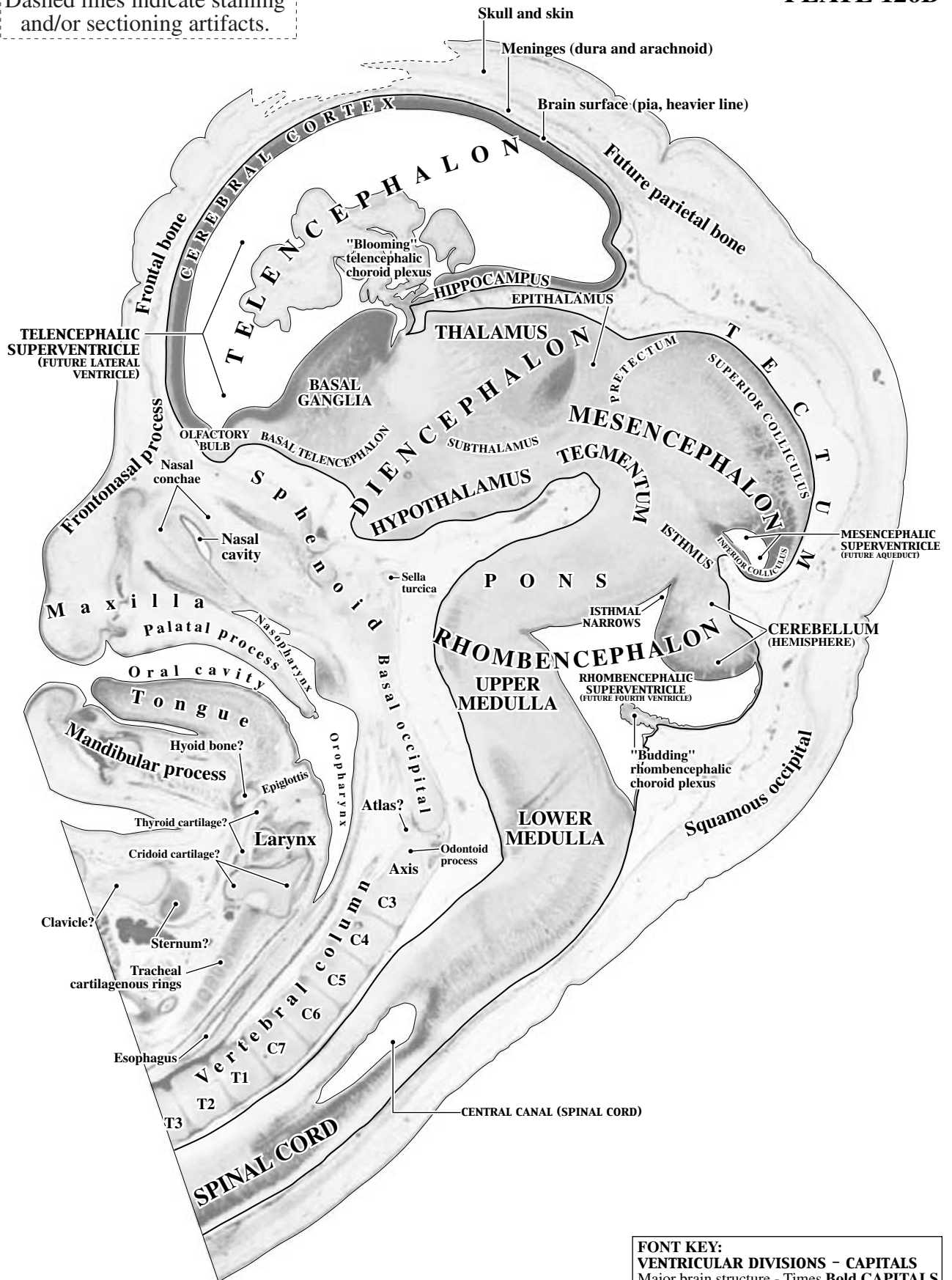
2 mm





Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

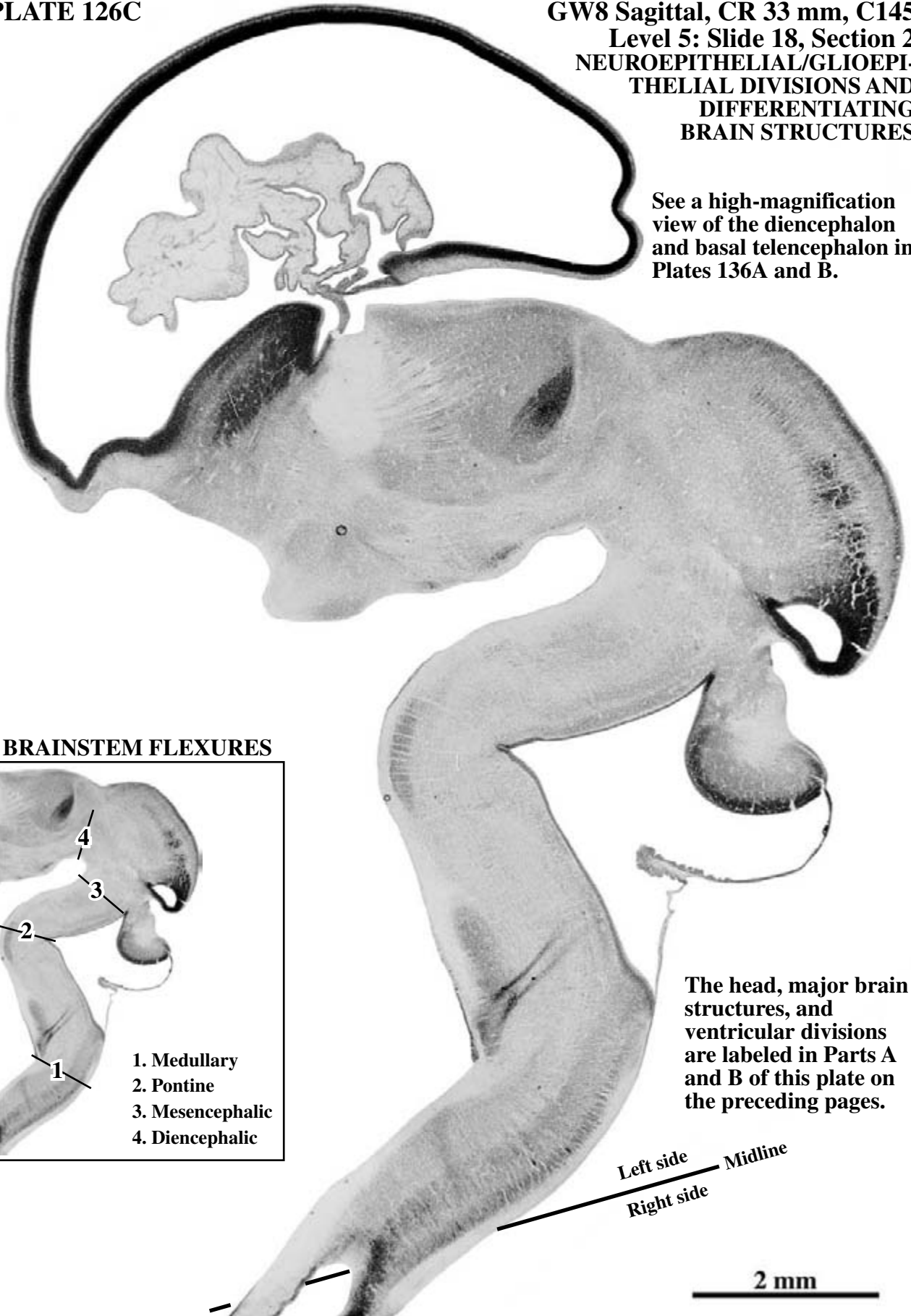
Dashed lines indicate staining and/or sectioning artifacts.



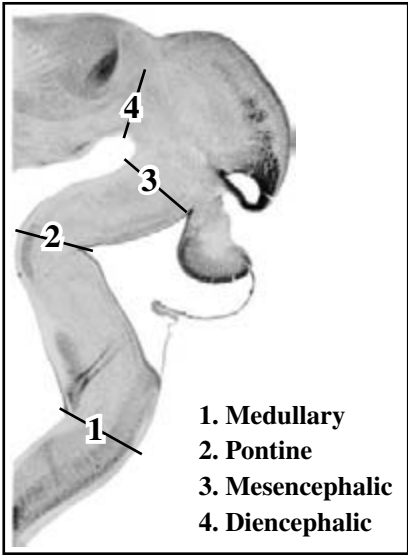
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Major brain structure - Times **Bold CAPITALS**
All other structures - Times Roman or **Bold**

**GW8 Sagittal, CR 33 mm, C145
Level 5: Slide 18, Section 2
NEUROEPITHELIAL/GLIOEPI-
THELIAL DIVISIONS AND
DIFFERENTIATING
BRAIN STRUCTURES**

See a high-magnification view of the diencephalon and basal telencephalon in Plates 136A and B.



BRAINSTEM FLEXURES

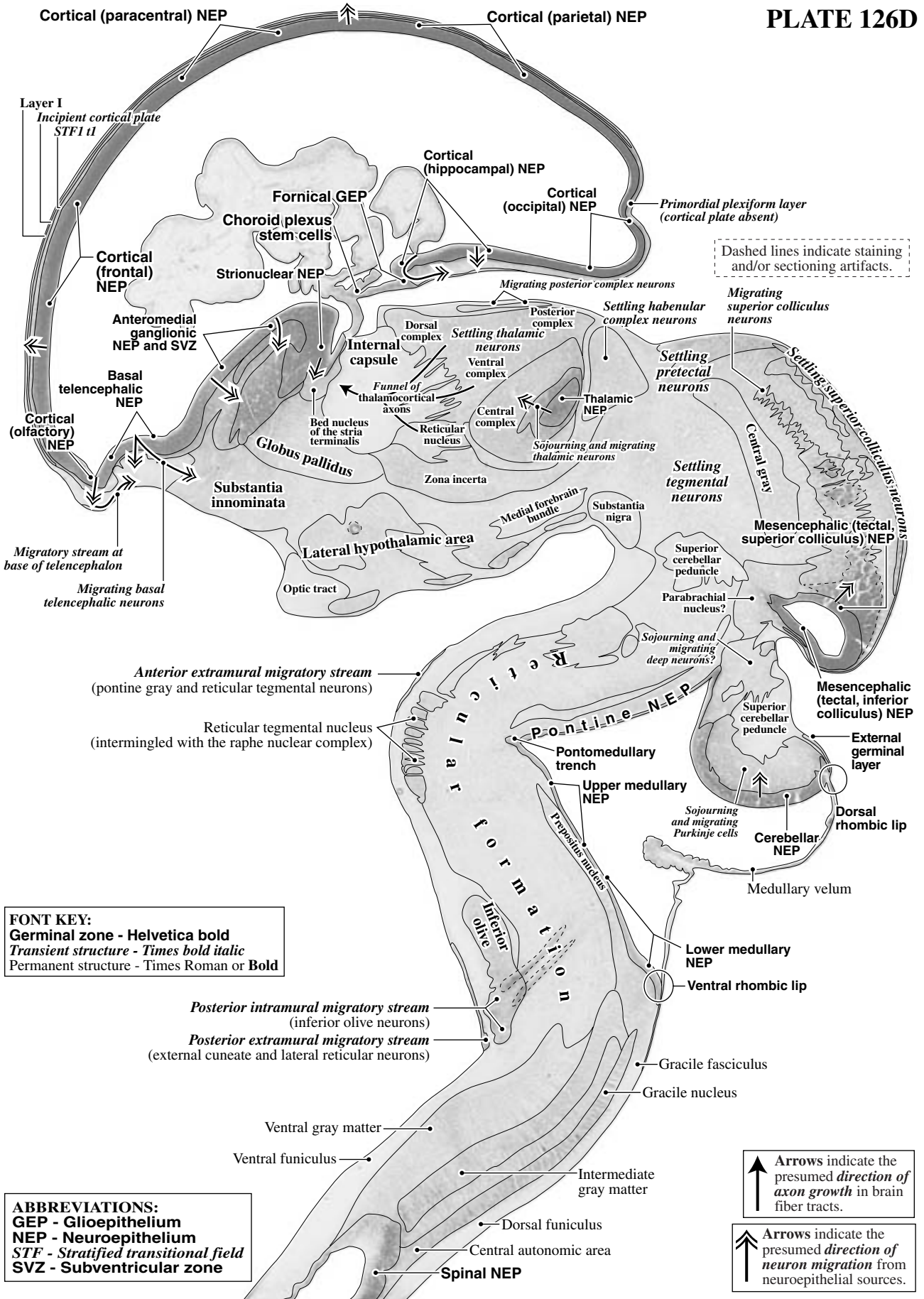


- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

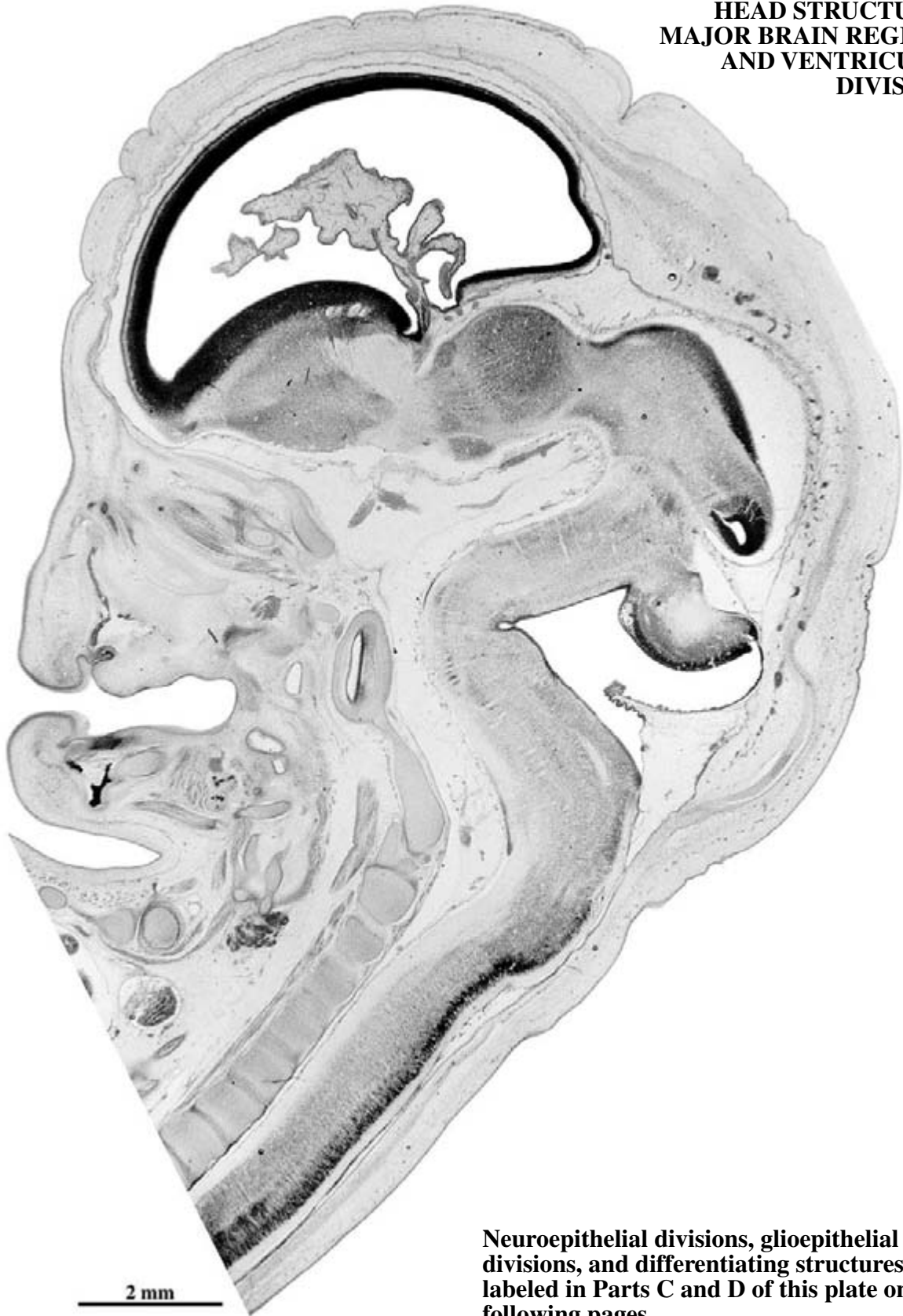
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

Left side
Right side
Midline

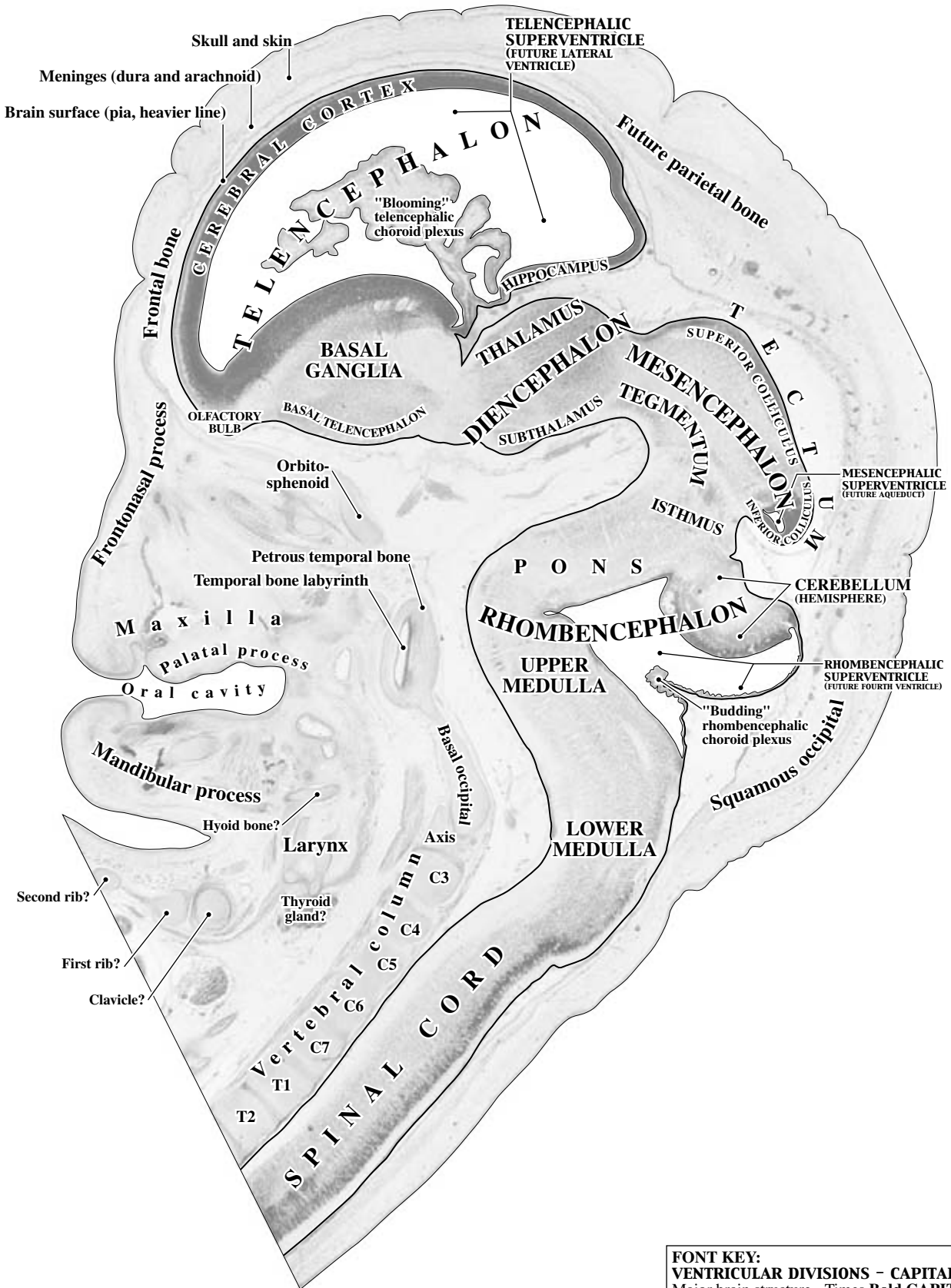
2 mm



GW8 Sagittal, CR 33 mm, C145
Level 6: Slide 16, Section 2
Left side of brain
HEAD STRUCTURES,
MAJOR BRAIN REGIONS,
AND VENTRICULAR
DIVISIONS



Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



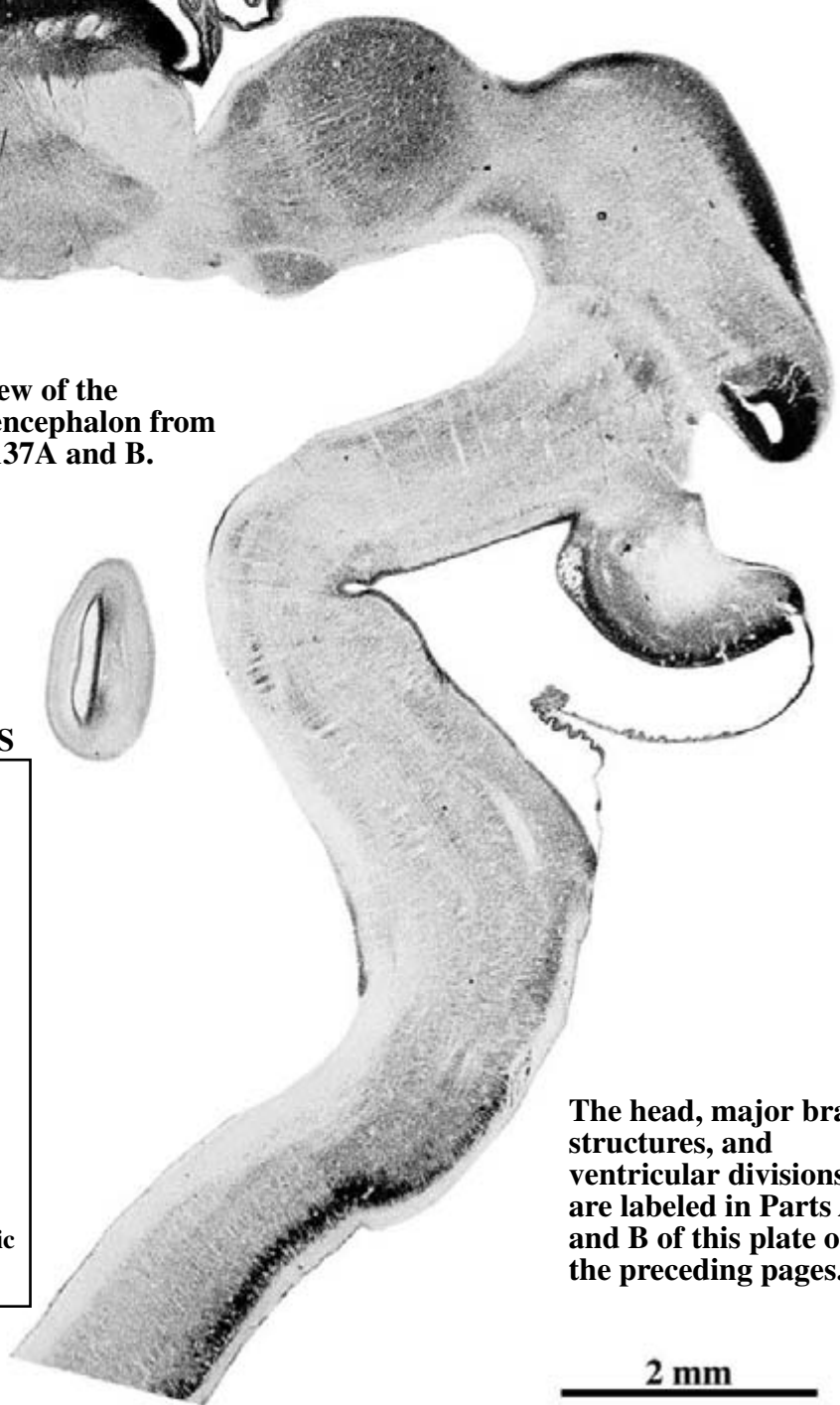
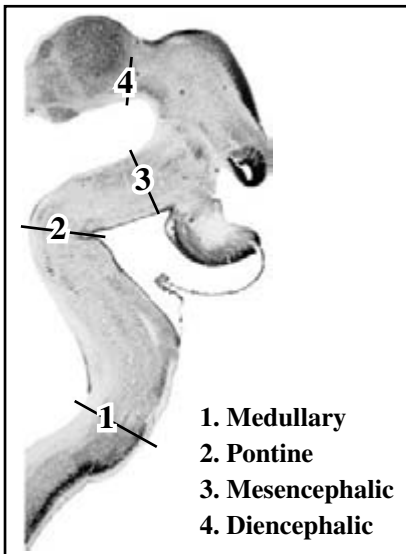
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

PLATE 127C

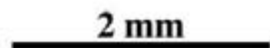
GW8 Sagittal, CR 33 mm, C145
Level 6: Slide 16, Section 2
Left side of brain
NEUROEPITHELIAL/
GLIOEPITHELIAL
DIVISIONS AND
DIFFERENTIATING
BRAIN STRUCTURES

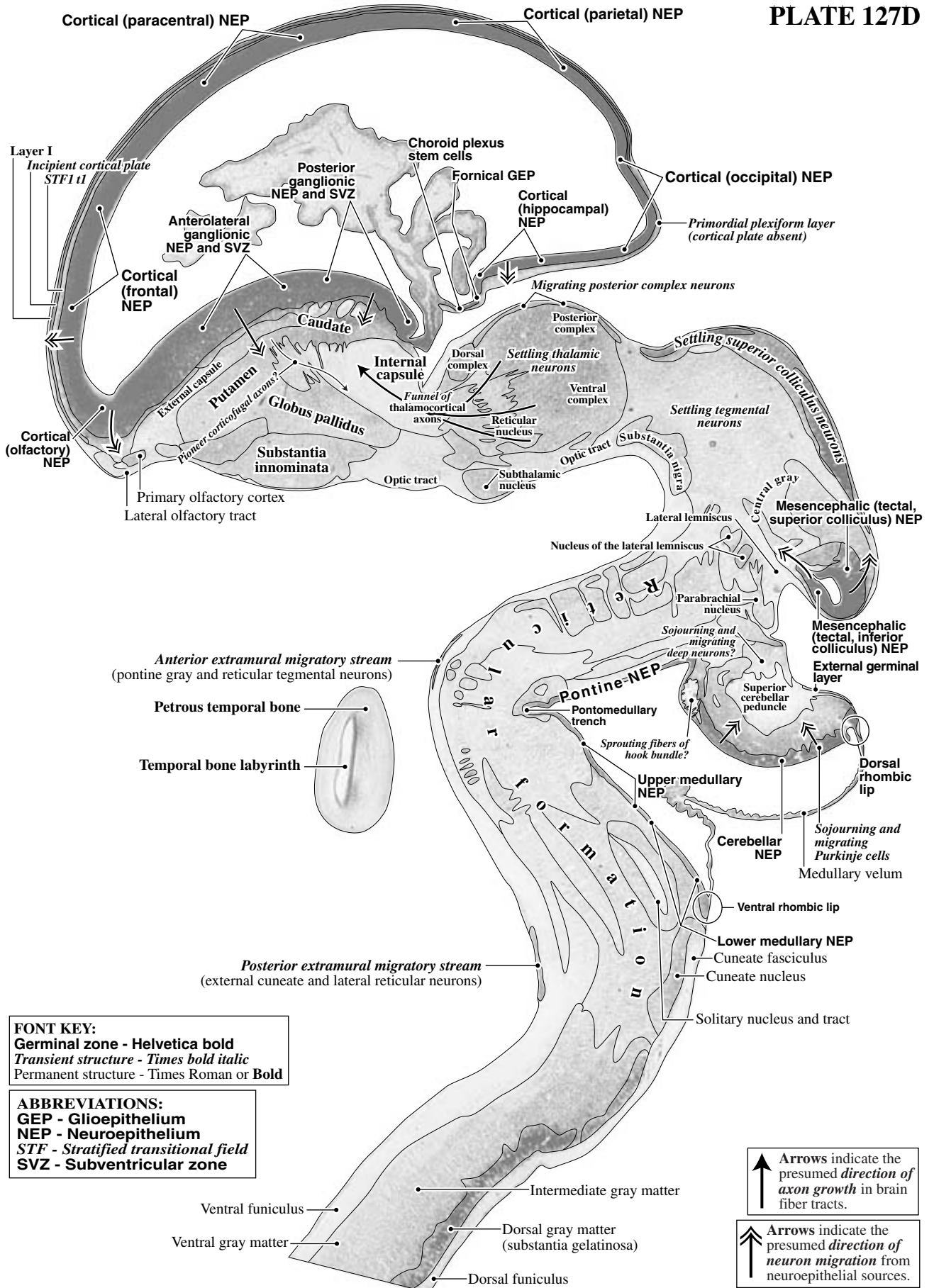
See a high-magnification view of the diencephalon and basal telencephalon from slide 17 section 2 in Plates 137A and B.

BRAINSTEM FLEXURES



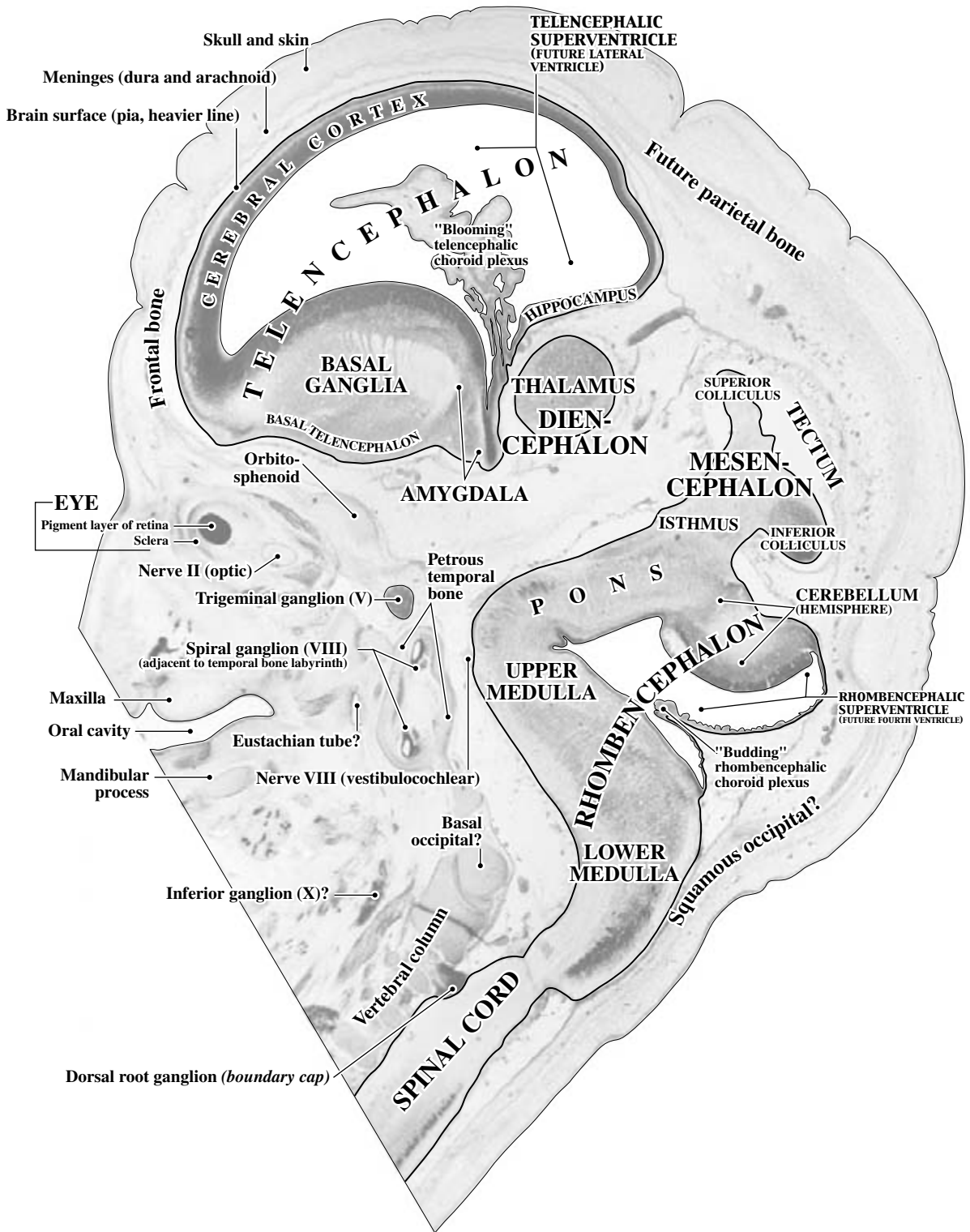
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.







Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



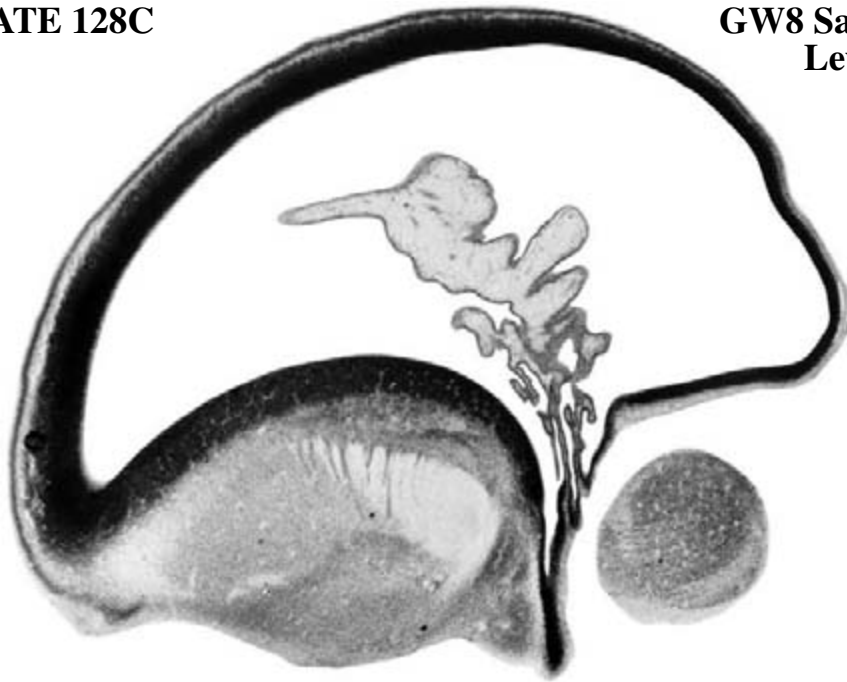
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

PLATE 128C

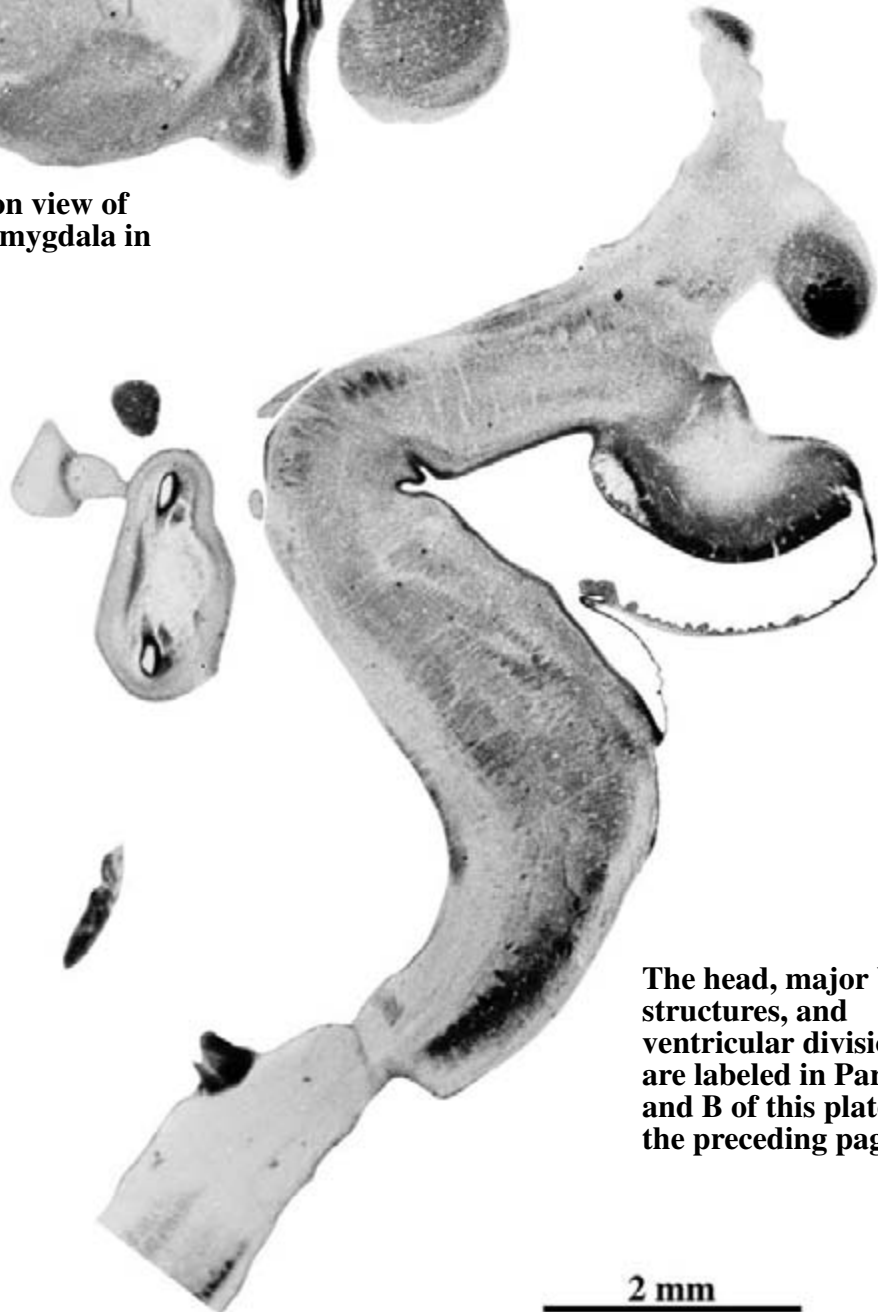
GW8 Sagittal, CR 33 mm, C145

Level 7: Slide 15, Section 1

Left side of brain
 NEUROEPITHELIAL/
 GLIOEPITHELIAL
 DIVISIONS AND
 DIFFERENTIATING
 BRAIN STRUCTURES

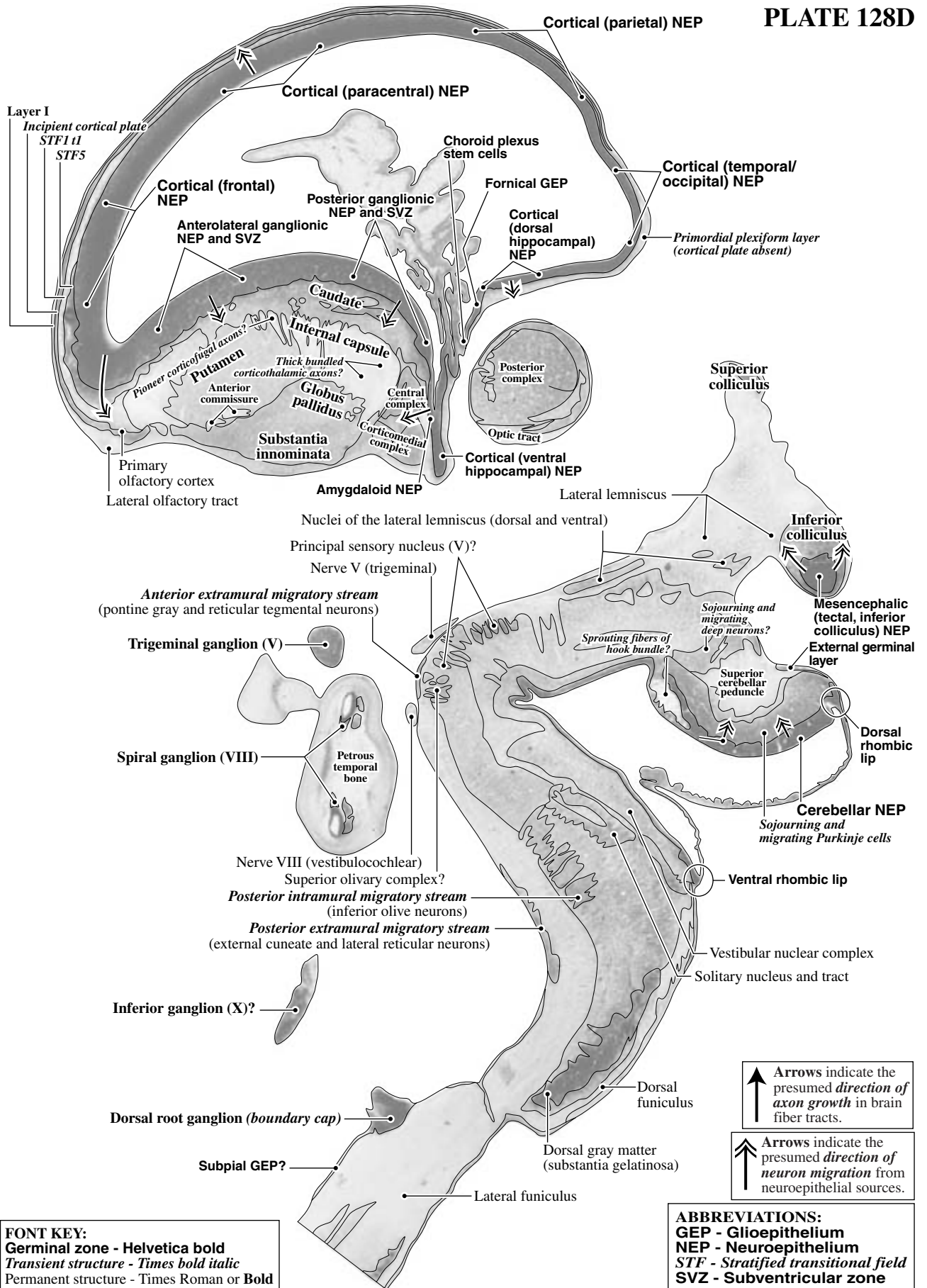


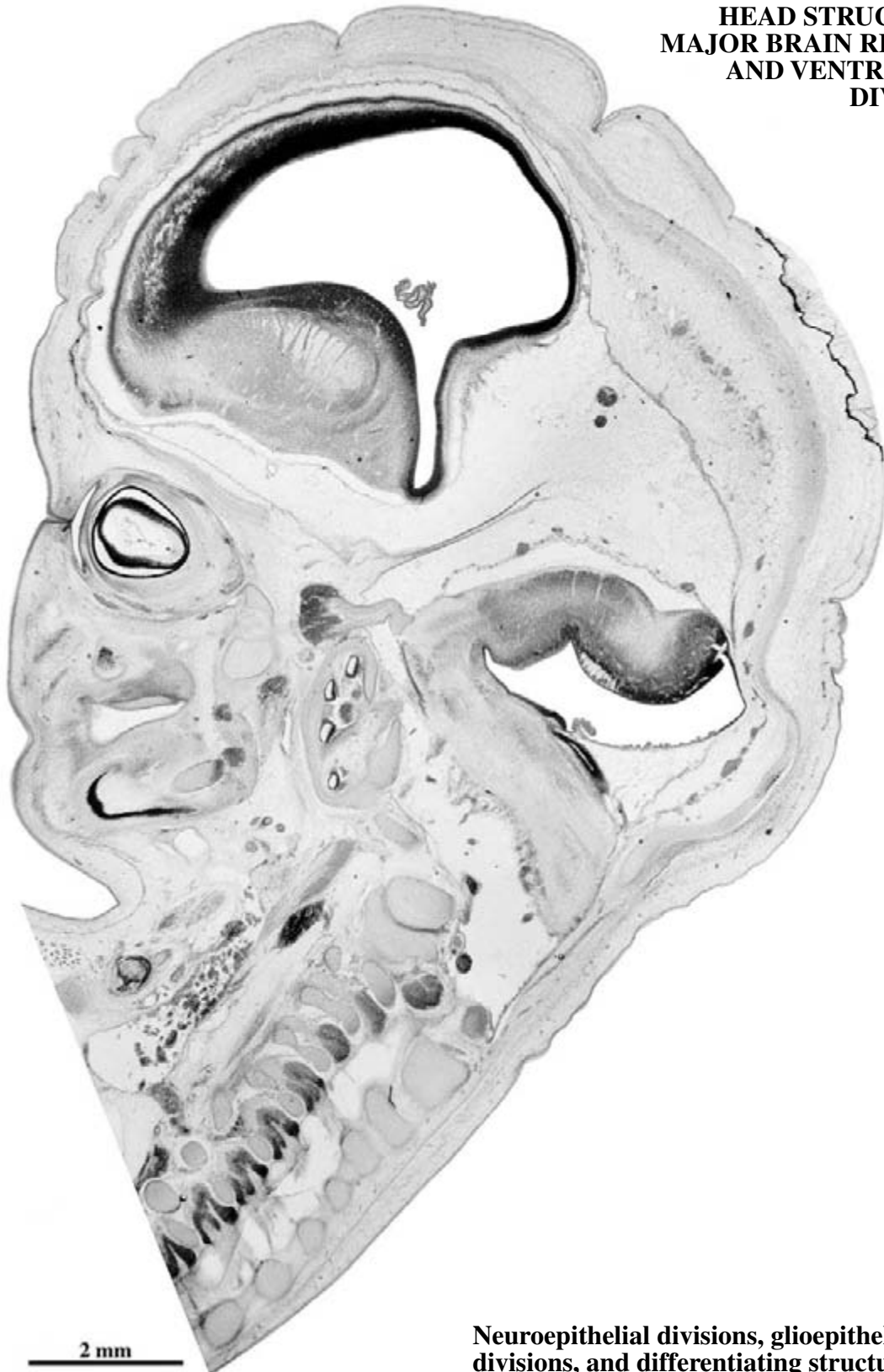
See a high-magnification view of the basal ganglia and amygdala in Plates 132A and B.



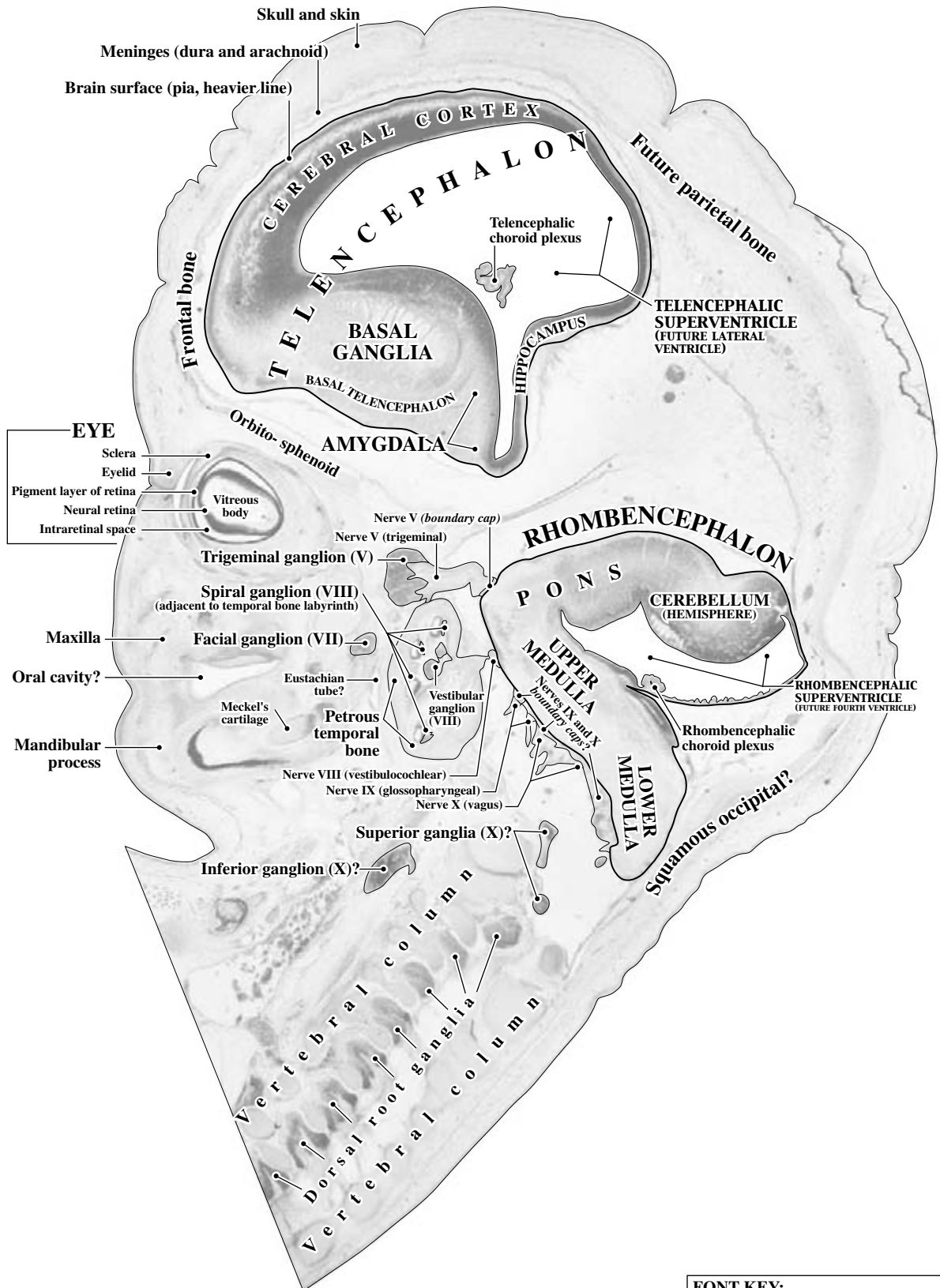
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

2 mm

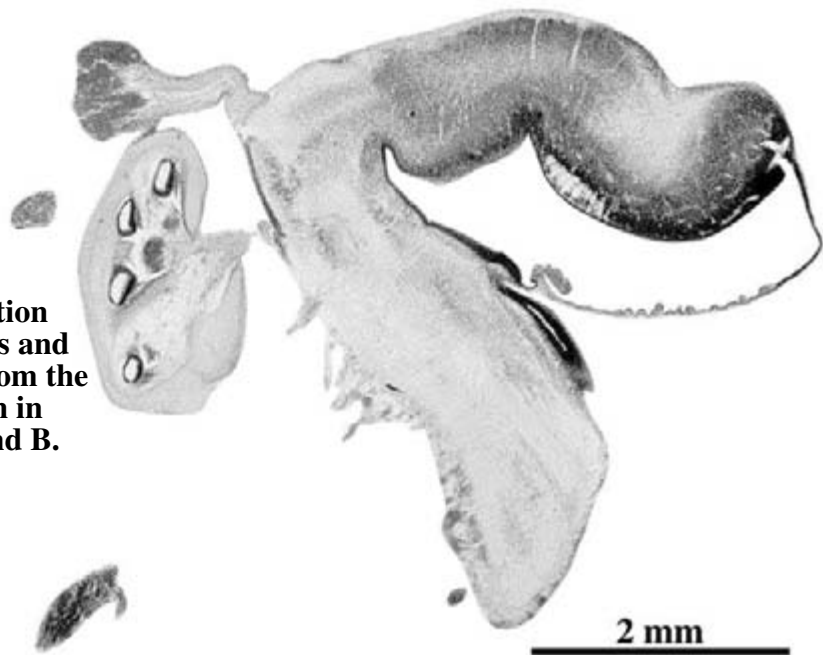
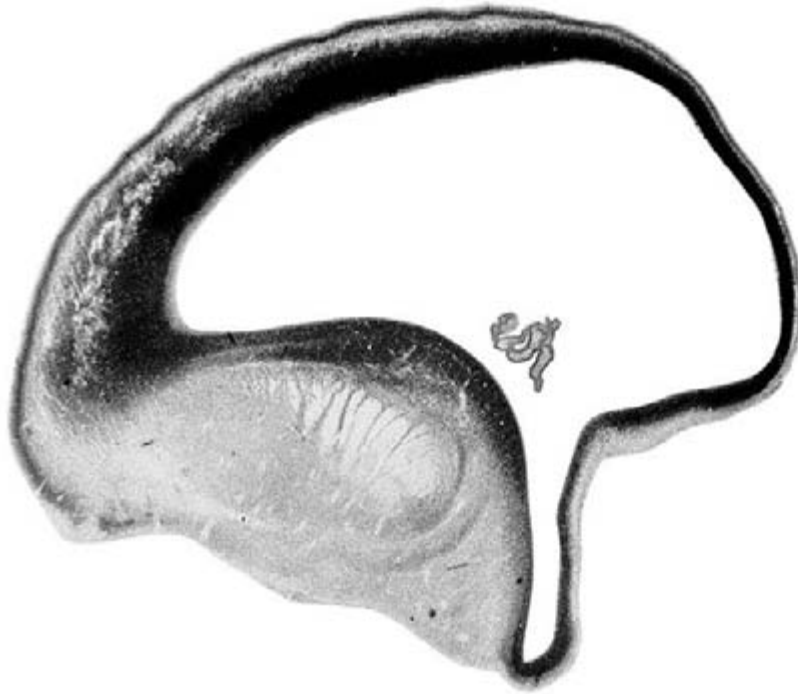




Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.

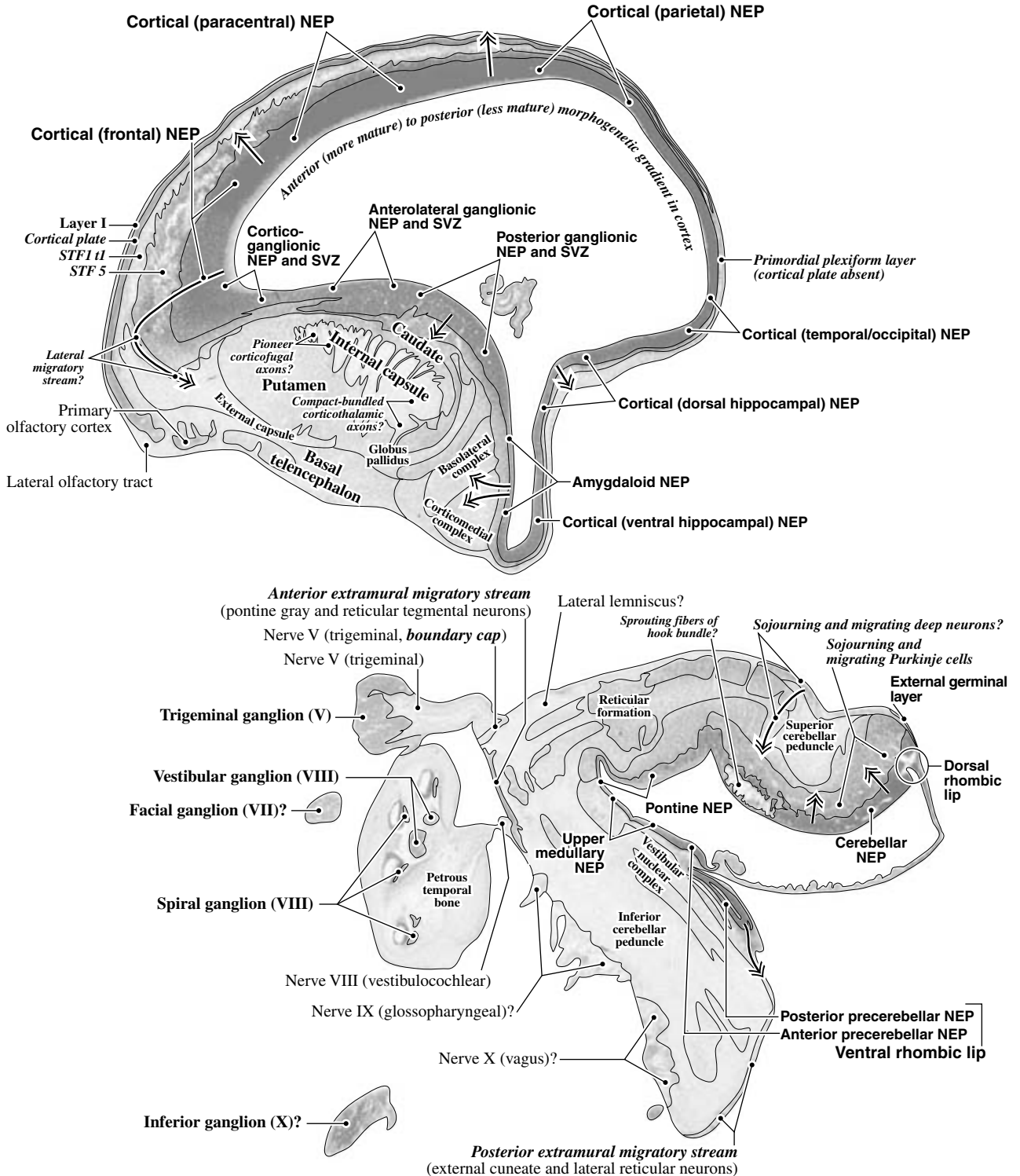


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



See a high-magnification view of the basal pons and peripheral ganglia from the right side of the brain in Plates 146 to 147A and B.

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

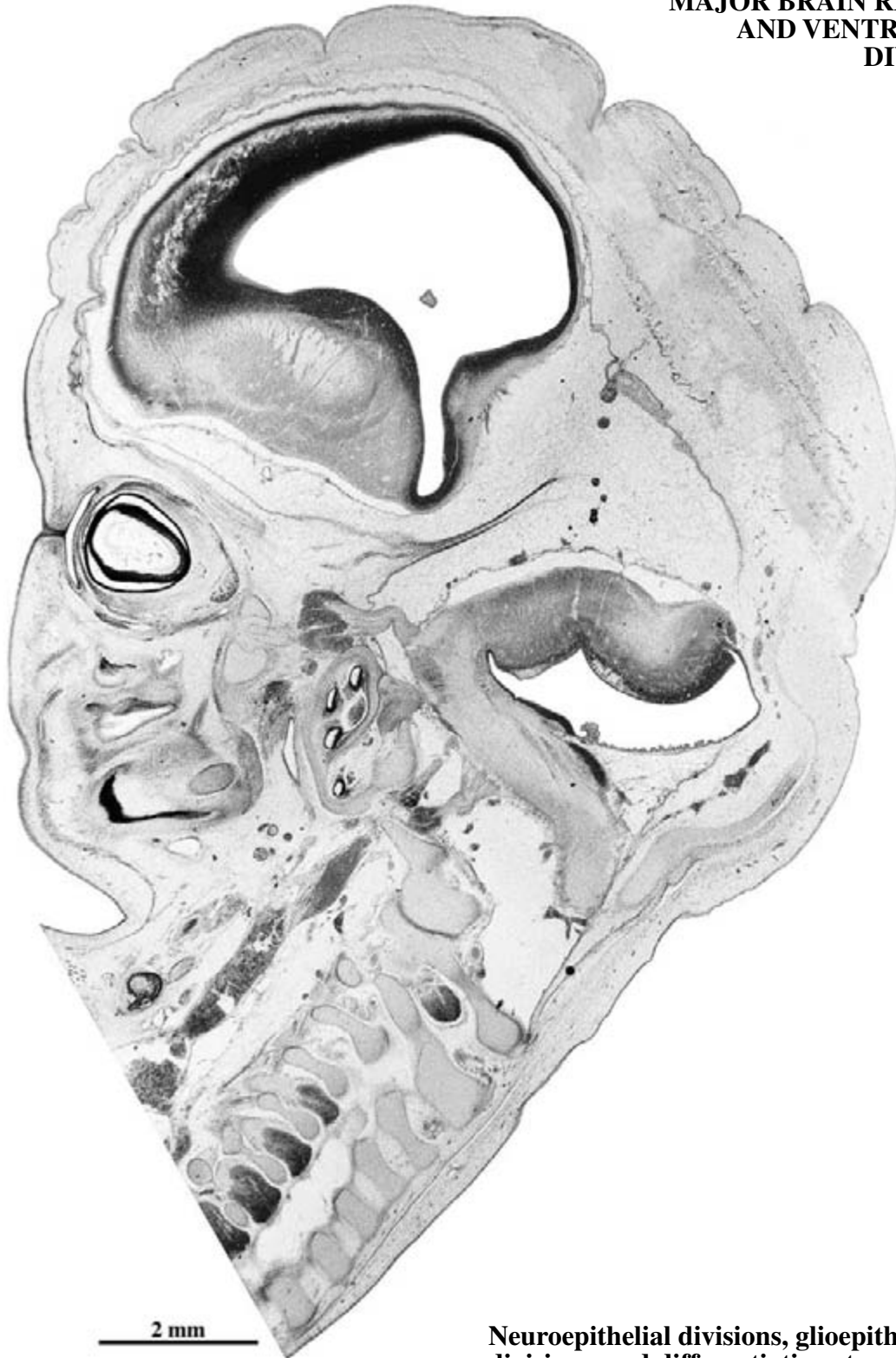


FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

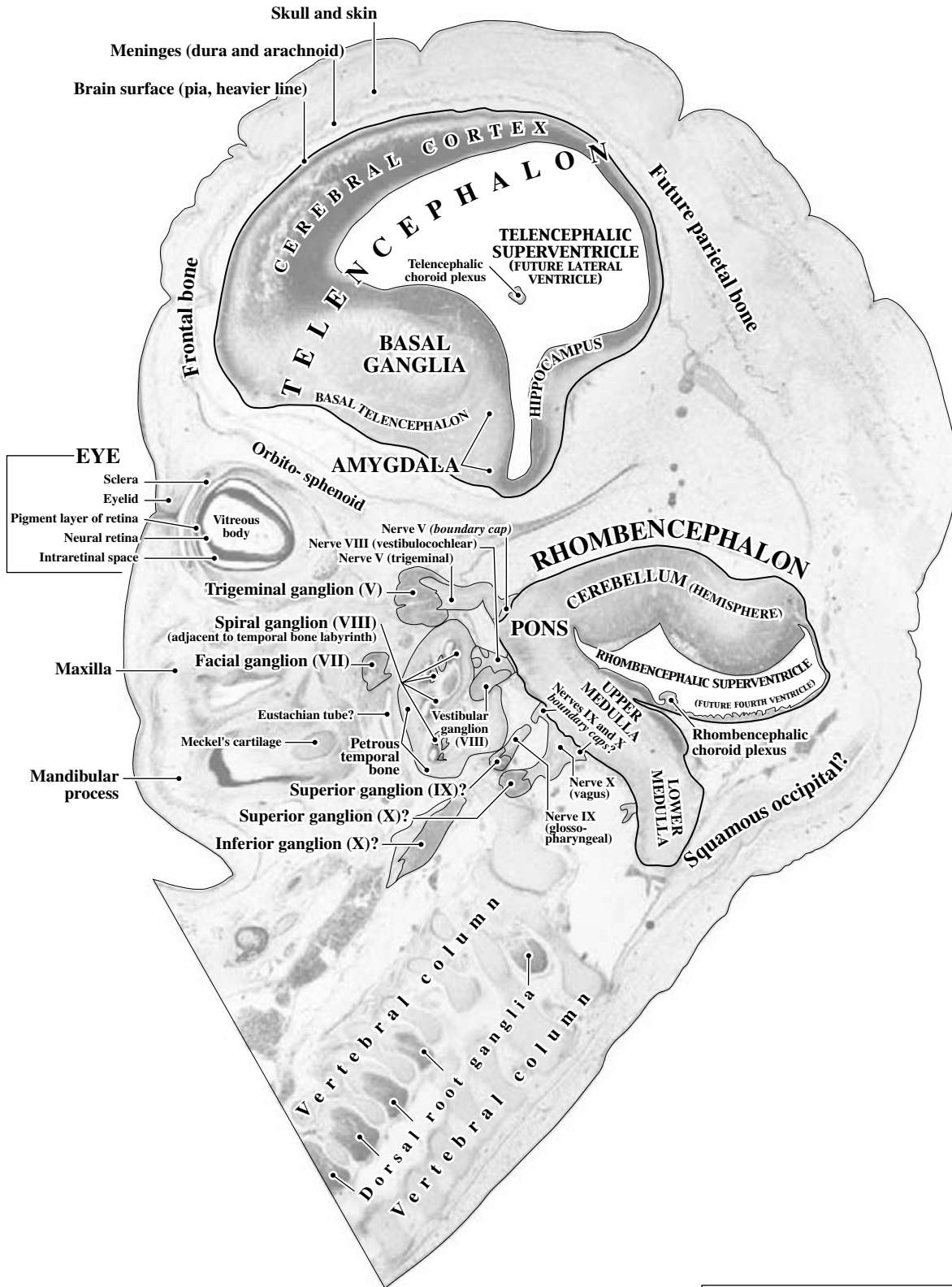
ABBREVIATIONS:
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

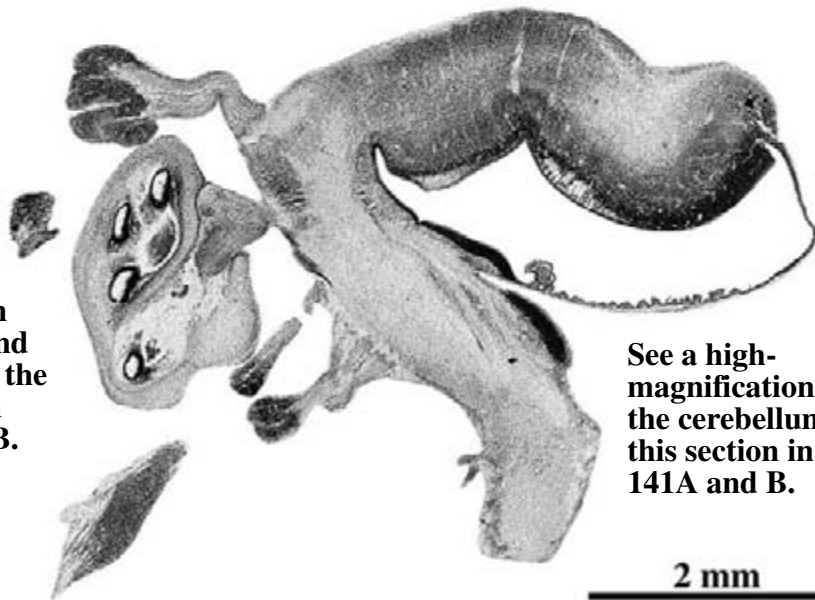
**GW8 Sagittal, CR 33 mm, C145
Level 9: Slide 12, Section 4
Left side of brain
HEAD STRUCTURES,
MAJOR BRAIN REGIONS,
AND VENTRICULAR
DIVISIONS**



Neuroepithelial divisions, gliopithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

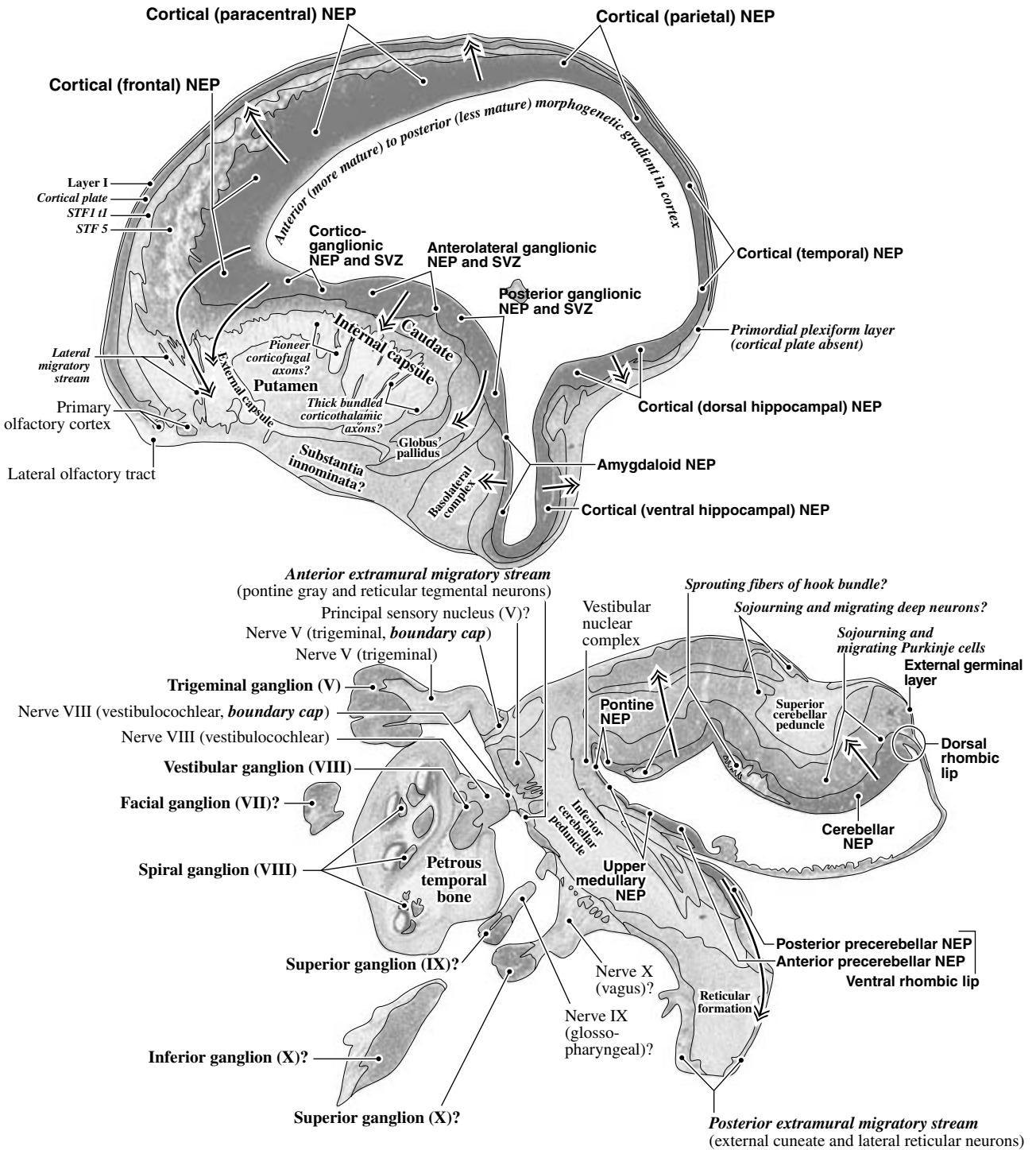


See a high-magnification view of the basal pons and peripheral ganglia from the right side of the brain in Plates 146 to 147A and B.

See a high-magnification view of the cerebellum from this section in Plates 141A and B.

2 mm

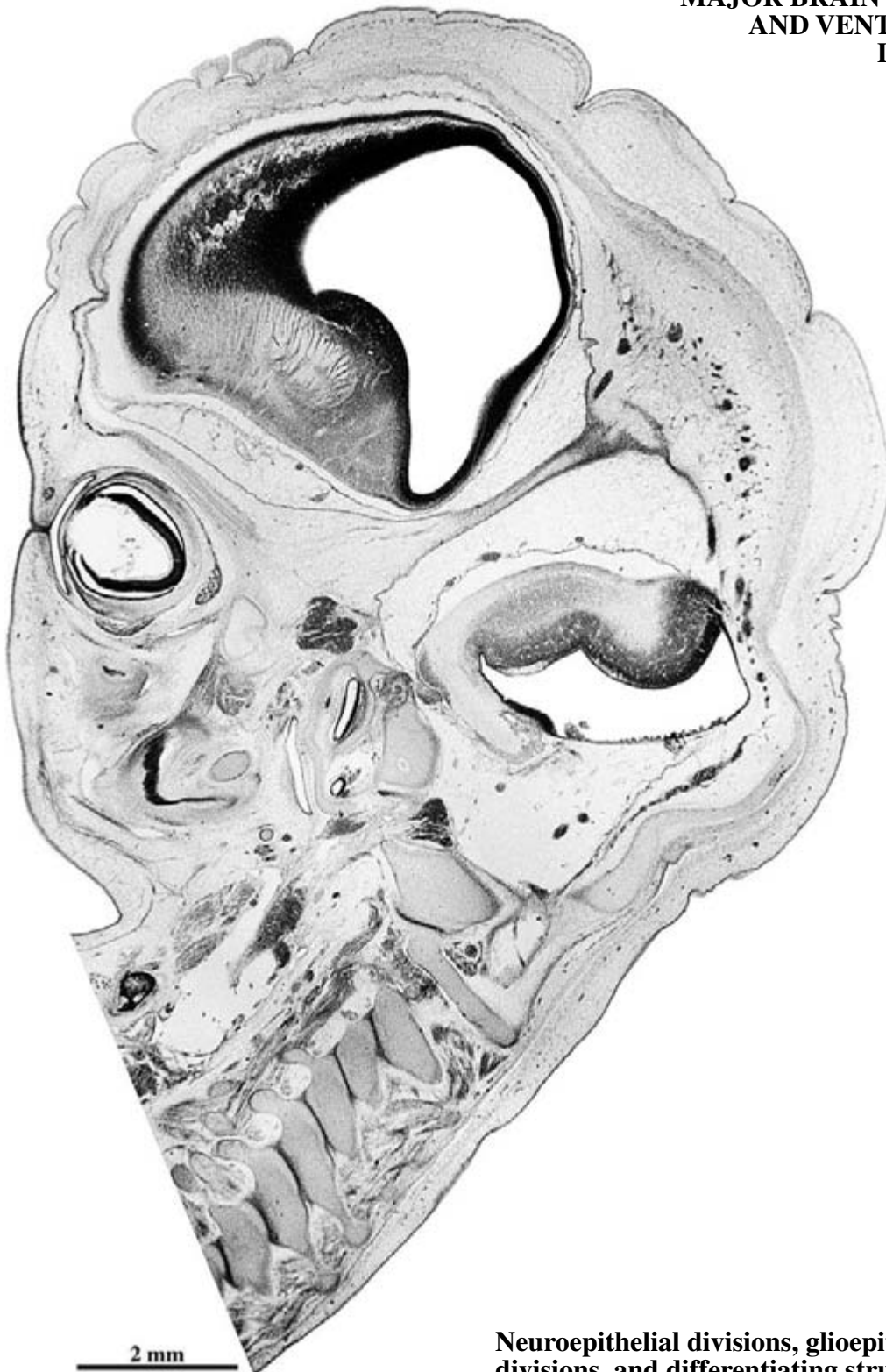
The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



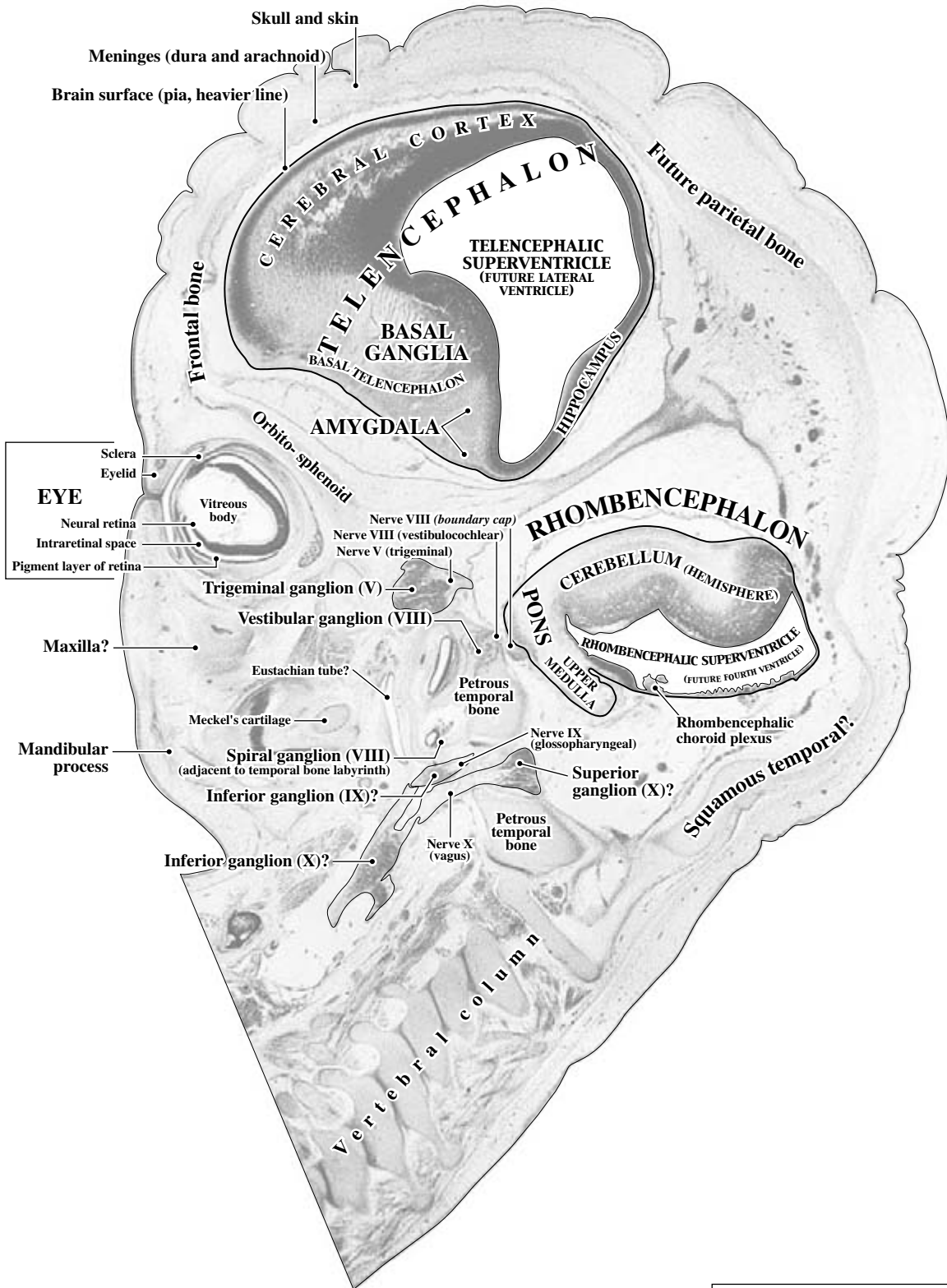
FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.



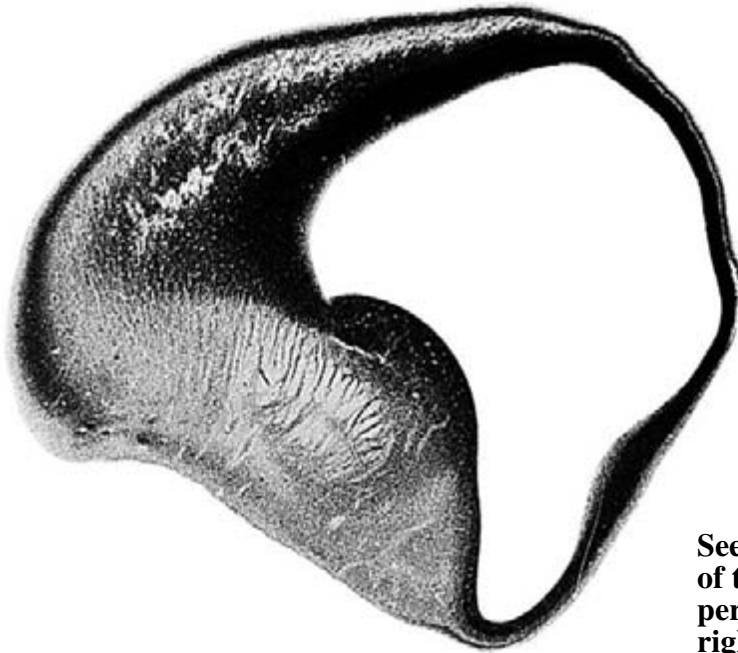
Neuroepithelial divisions, glioepithelial divisions, and differentiating structures are labeled in Parts C and D of this plate on the following pages.



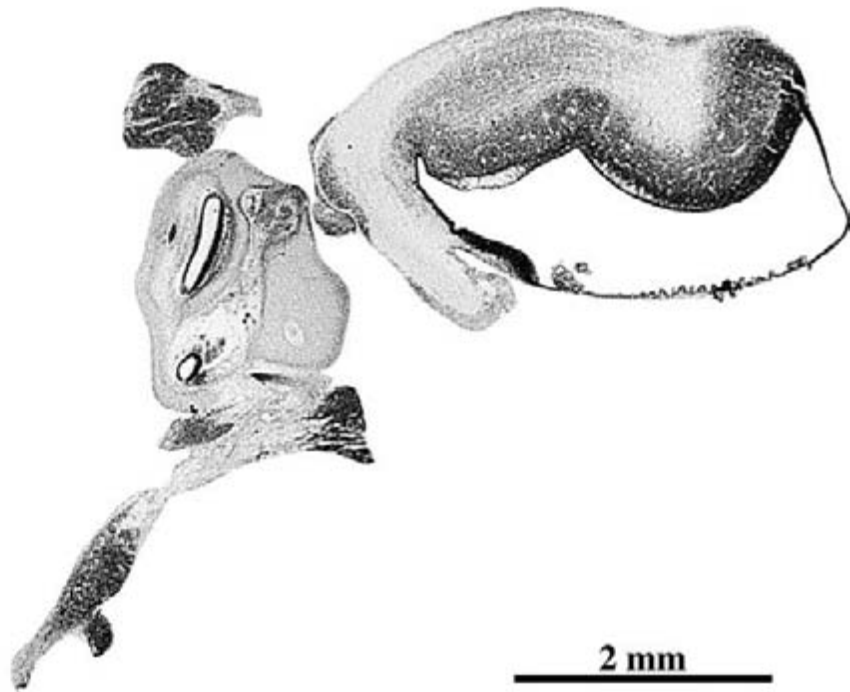
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

PLATE 131C

**GW8 Sagittal, CR 33 mm, C145
Level 10: Slide 11, Section 4
Left side of brain
NEUROEPITHELIAL/GLIOEPITHELIAL
DIVISIONS AND DIFFERENTIATING
BRAIN STRUCTURES**

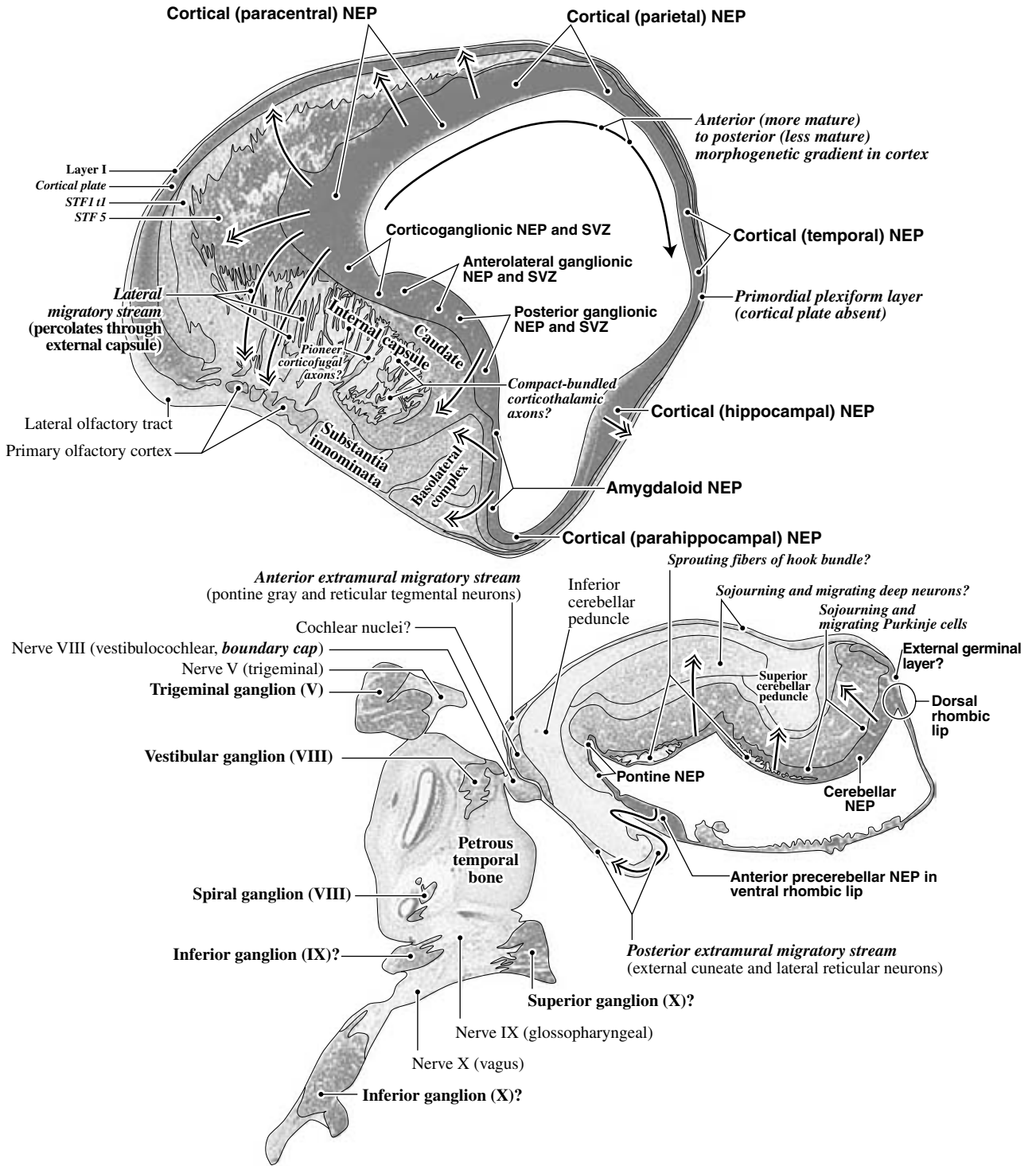


See a high-magnification view
of the basal pons and
peripheral ganglia from the
right side of the brain in Plates
146A and B to 147A and B.



2 mm

The head, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

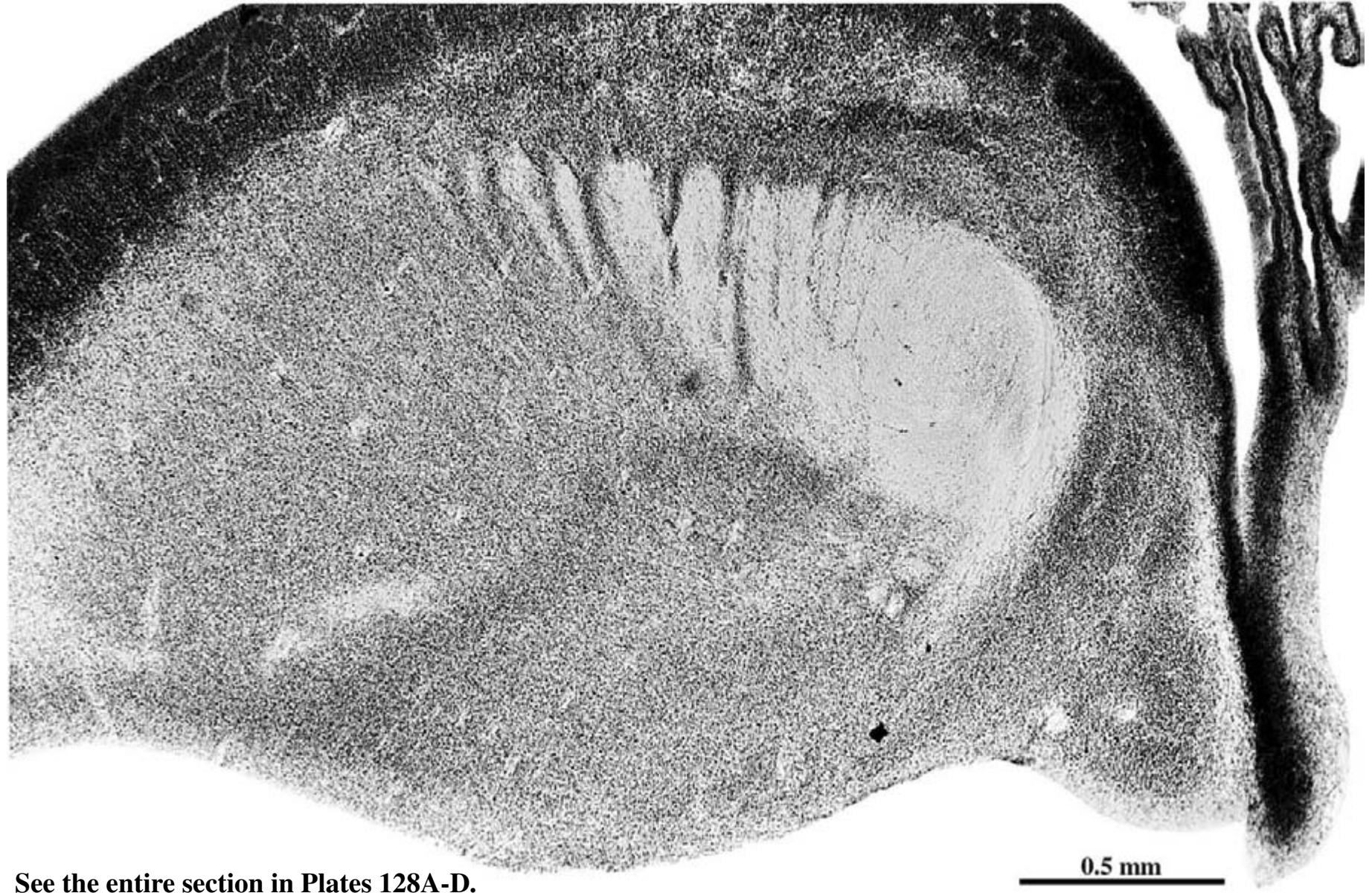
ABBREVIATIONS:
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 132A

**GW8 Sagittal, CR 33 mm, C145
Level 7: Slide 15, Section 1**

**BASAL GANGLIA, BASAL
TELENCEPHALON, AND AMYGDALA**



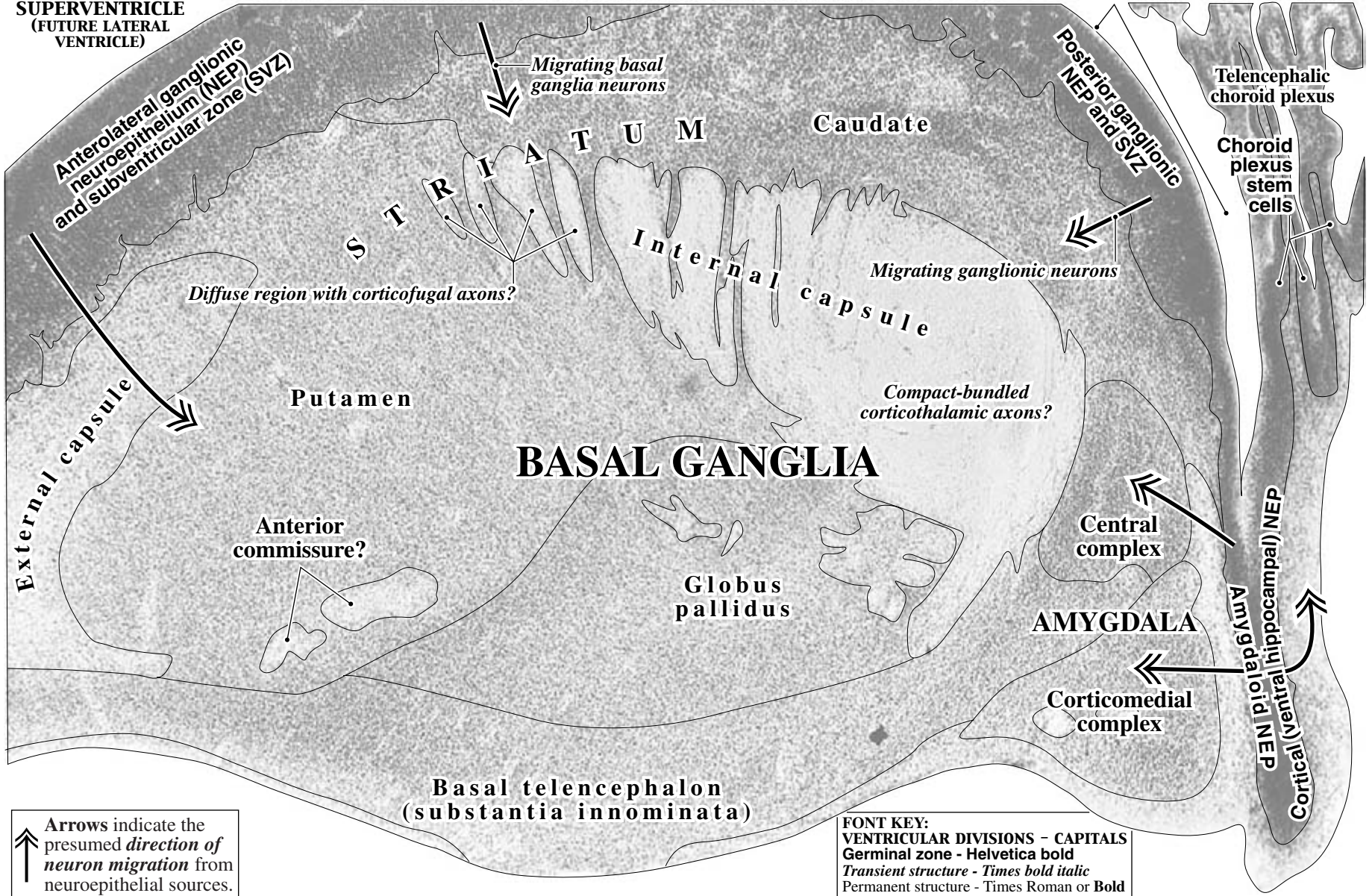
See the entire section in Plates 128A-D.

0.5 mm

PLATE 132B

**TELENCEPHALIC
SUPERVENTRICLE
(FUTURE LATERAL
VENTRICLE)**

**TELENCEPHALIC
SUPERVENTRICLE
(FUTURE LATERAL
VENTRICLE)**



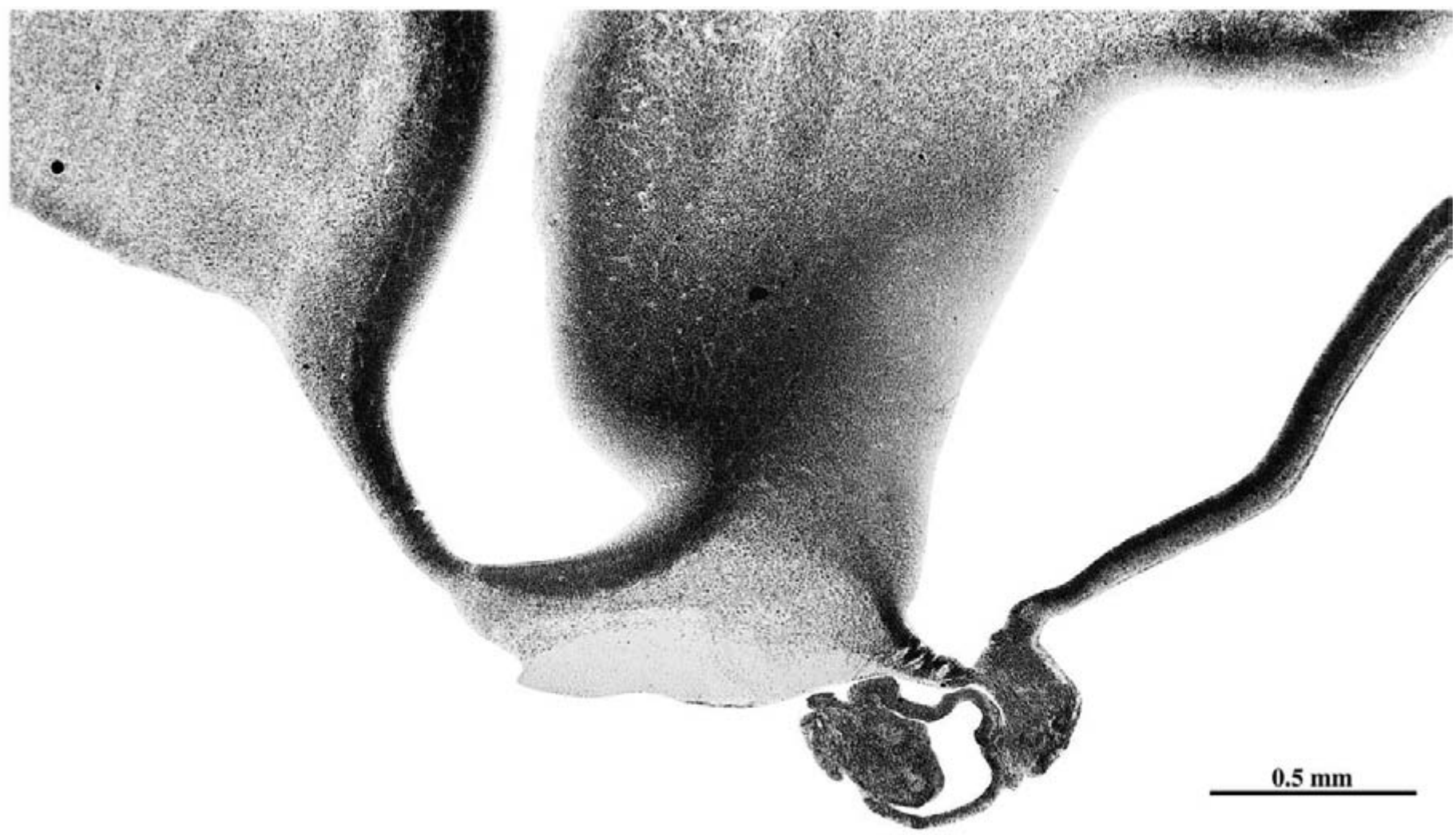
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 133A

**GW8 Sagittal, CR 33 mm, C145
Level 2: Slide 22, Section 2**

HYPOTHALAMUS AND PREOPTIC AREA



See the entire section in Plates 123A-D.

PLATE 133B

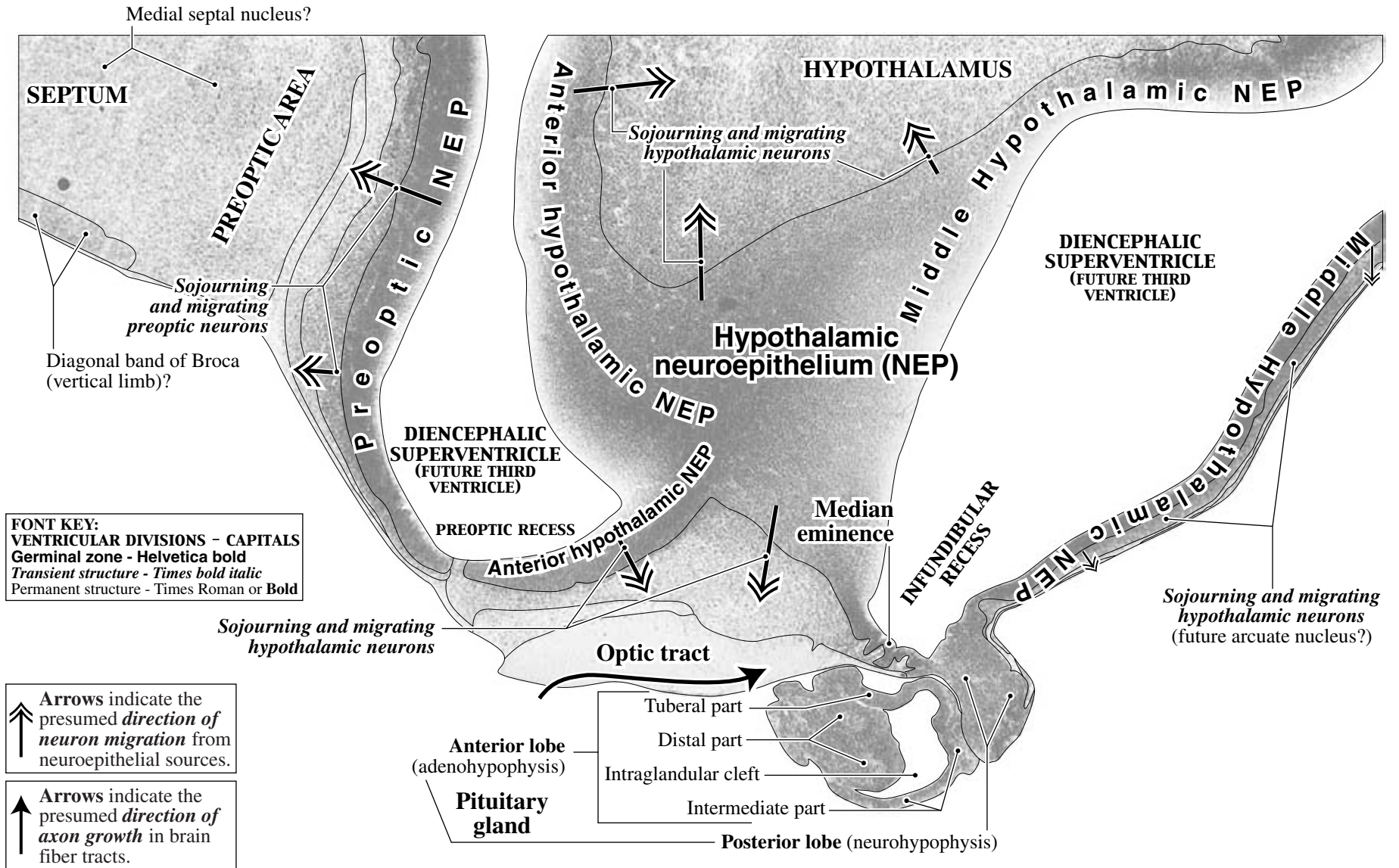
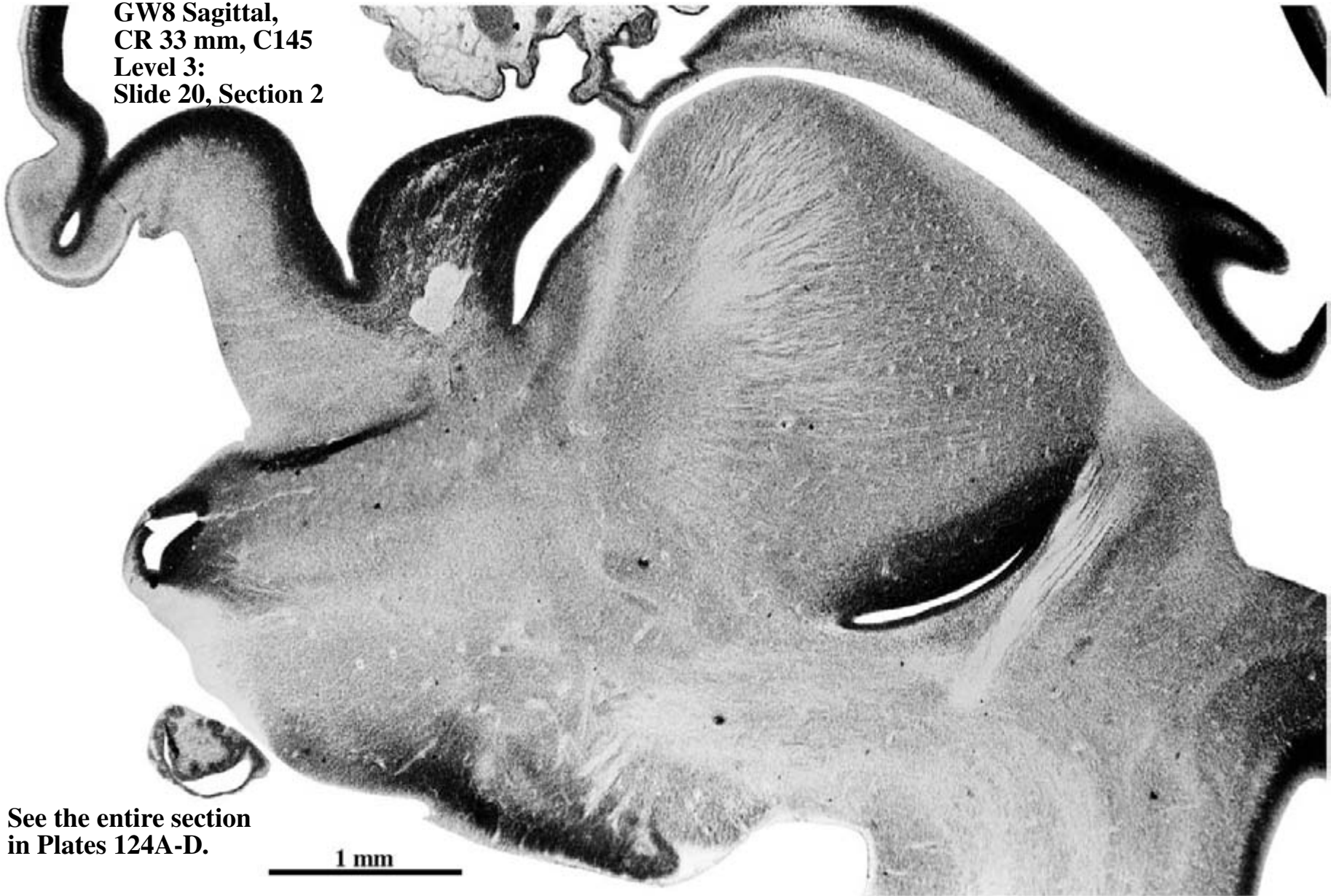


PLATE 134A

DIENCEPHALON AND BASAL TELENCEPHALON

**GW8 Sagittal,
CR 33 mm, C145
Level 3:
Slide 20, Section 2**



**See the entire section
in Plates 124A-D.**

1 mm

PLATE 134B

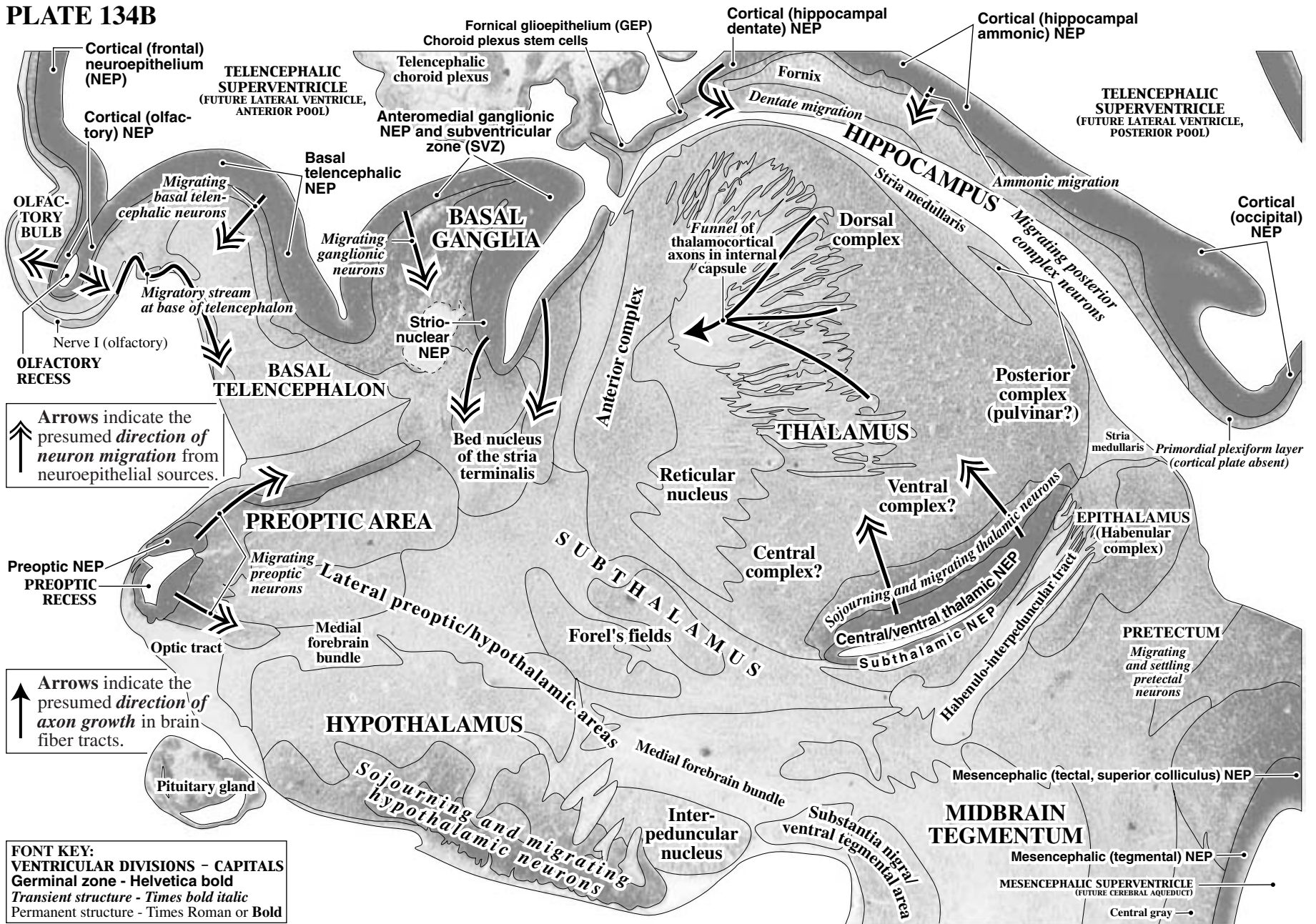
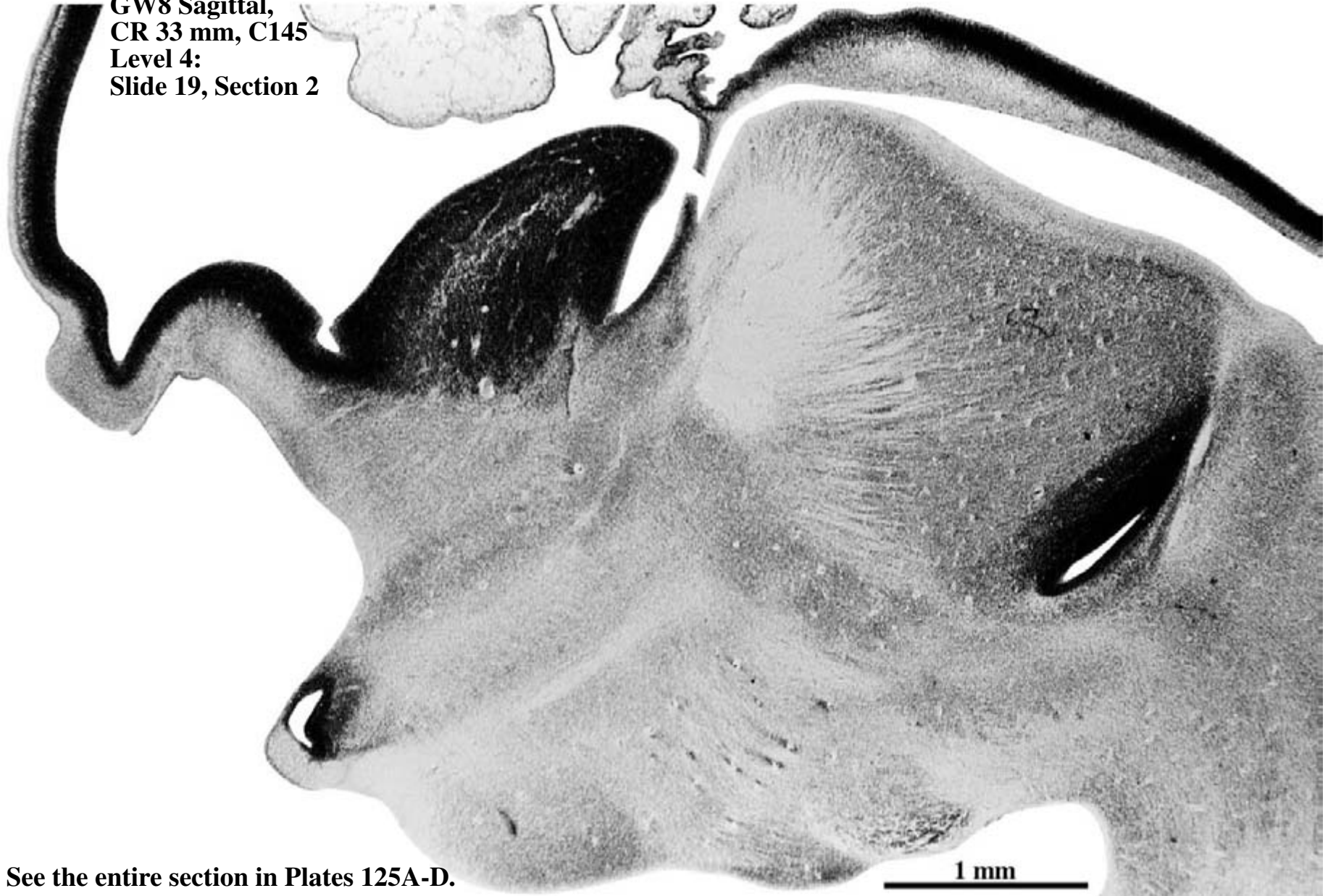


PLATE 135A

DIENCEPHALON AND BASAL TELENCEPHALON

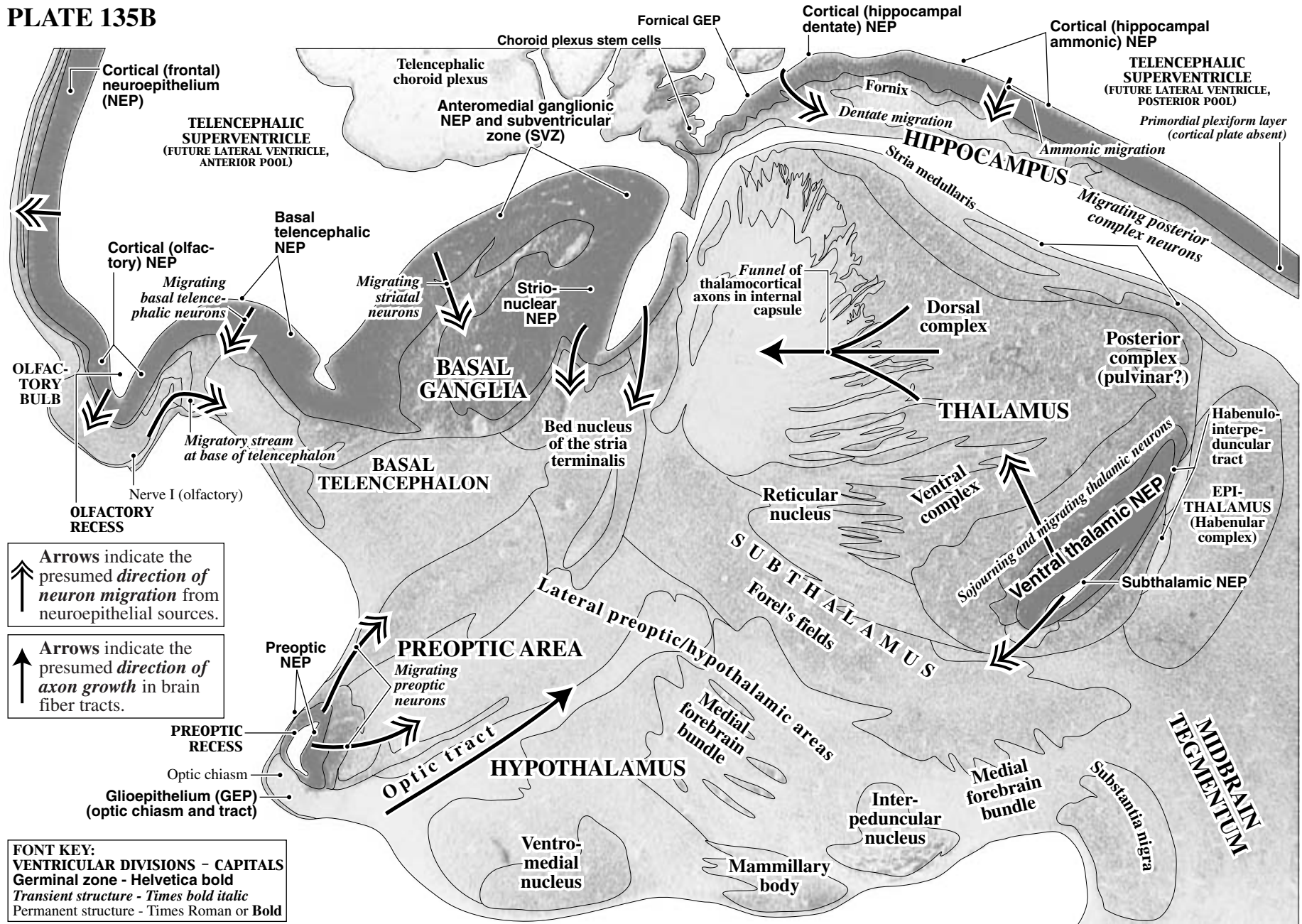
**GW8 Sagittal,
CR 33 mm, C145
Level 4:
Slide 19, Section 2**



See the entire section in Plates 125A-D.

1 mm

PLATE 135B



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 136A

**GW8 Sagittal, CR 33 mm, C145
Level 5: Slide 18, Section 2**

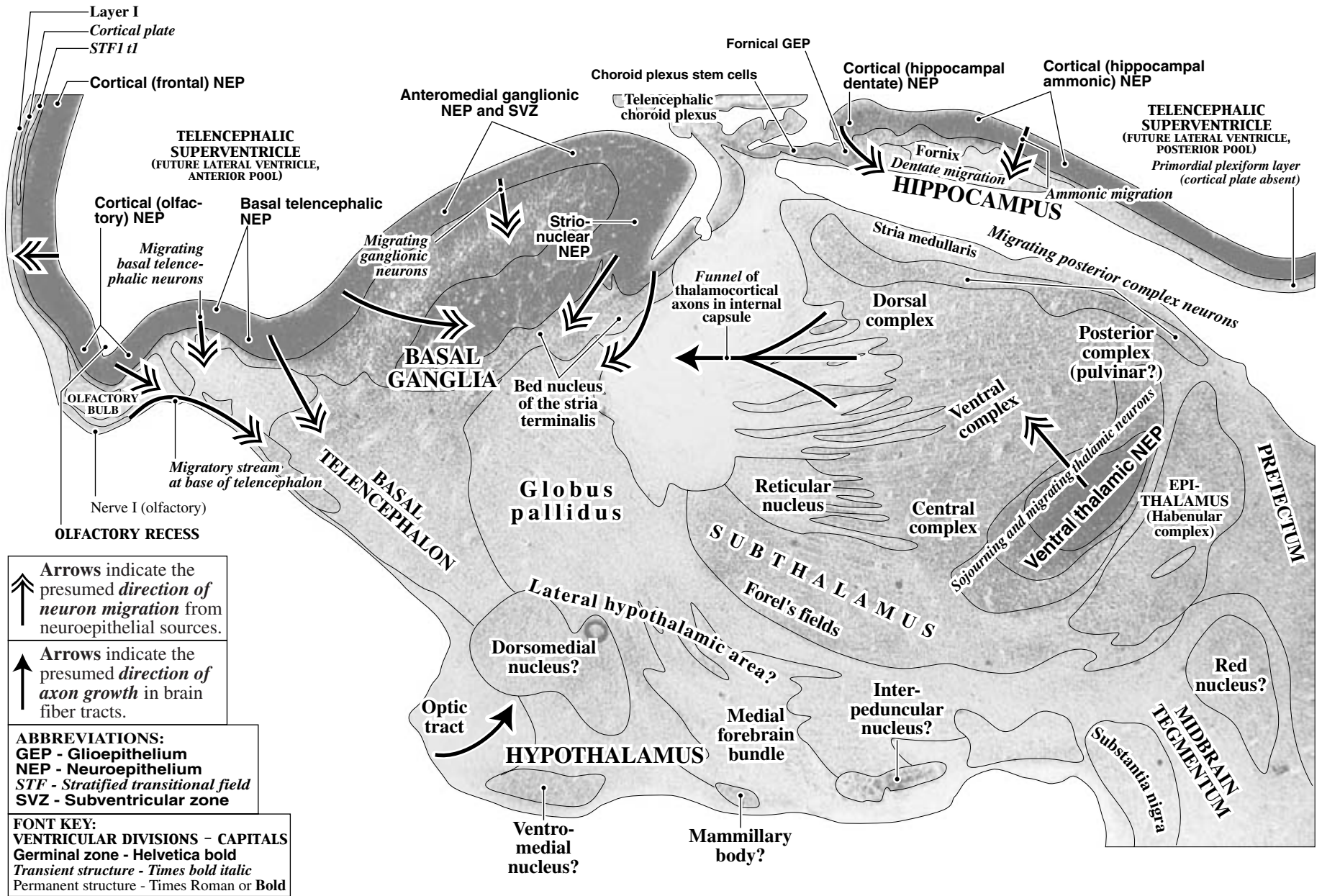
BASAL TELENCEPHALON AND DIENCEPHALON

354



See the entire section in Plates 126A-D.

PLATE 136B



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

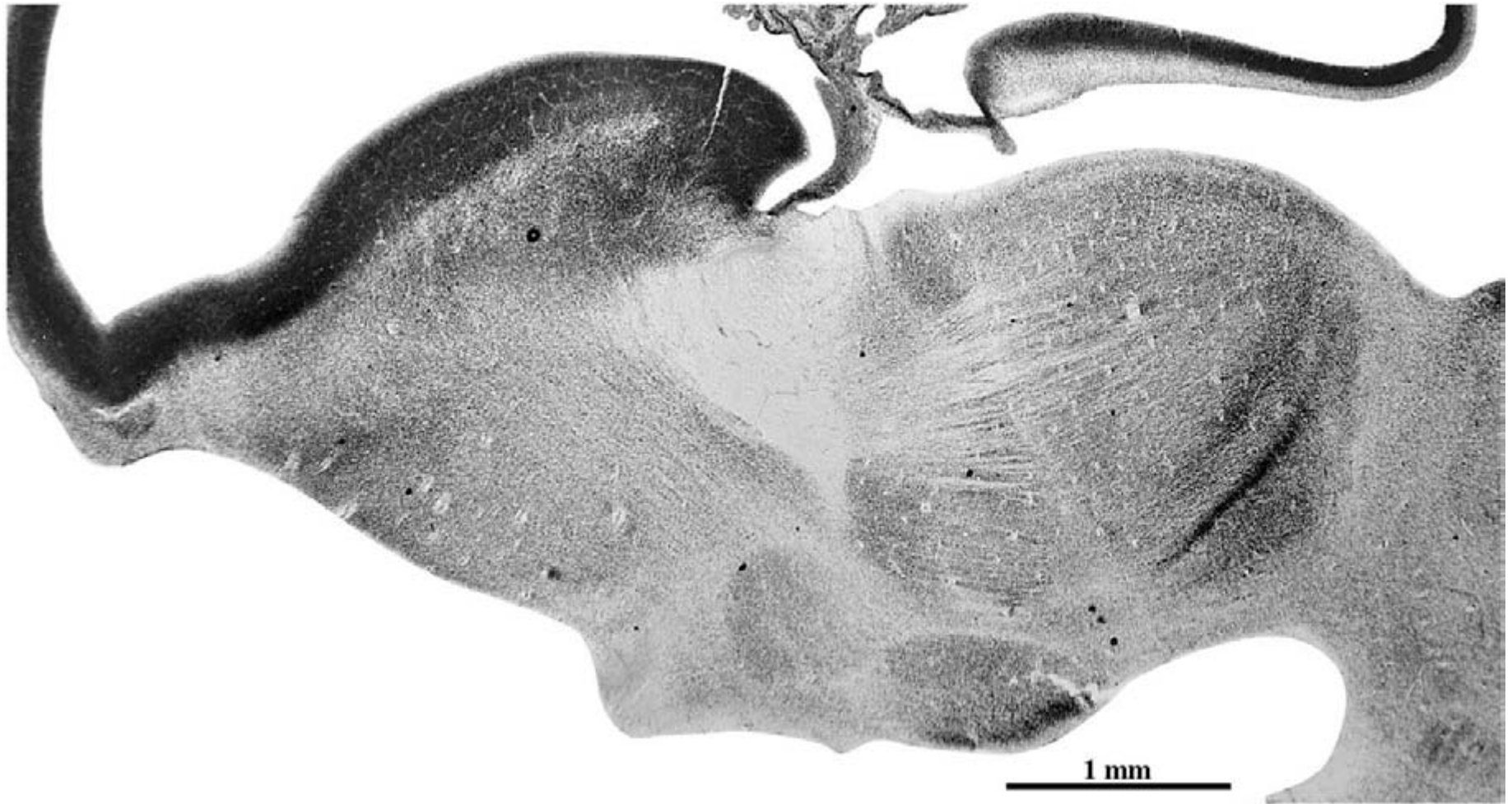
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
STF - Stratified transitional field
SVZ - Subventricular zone

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or **Bold**

PLATE 137A

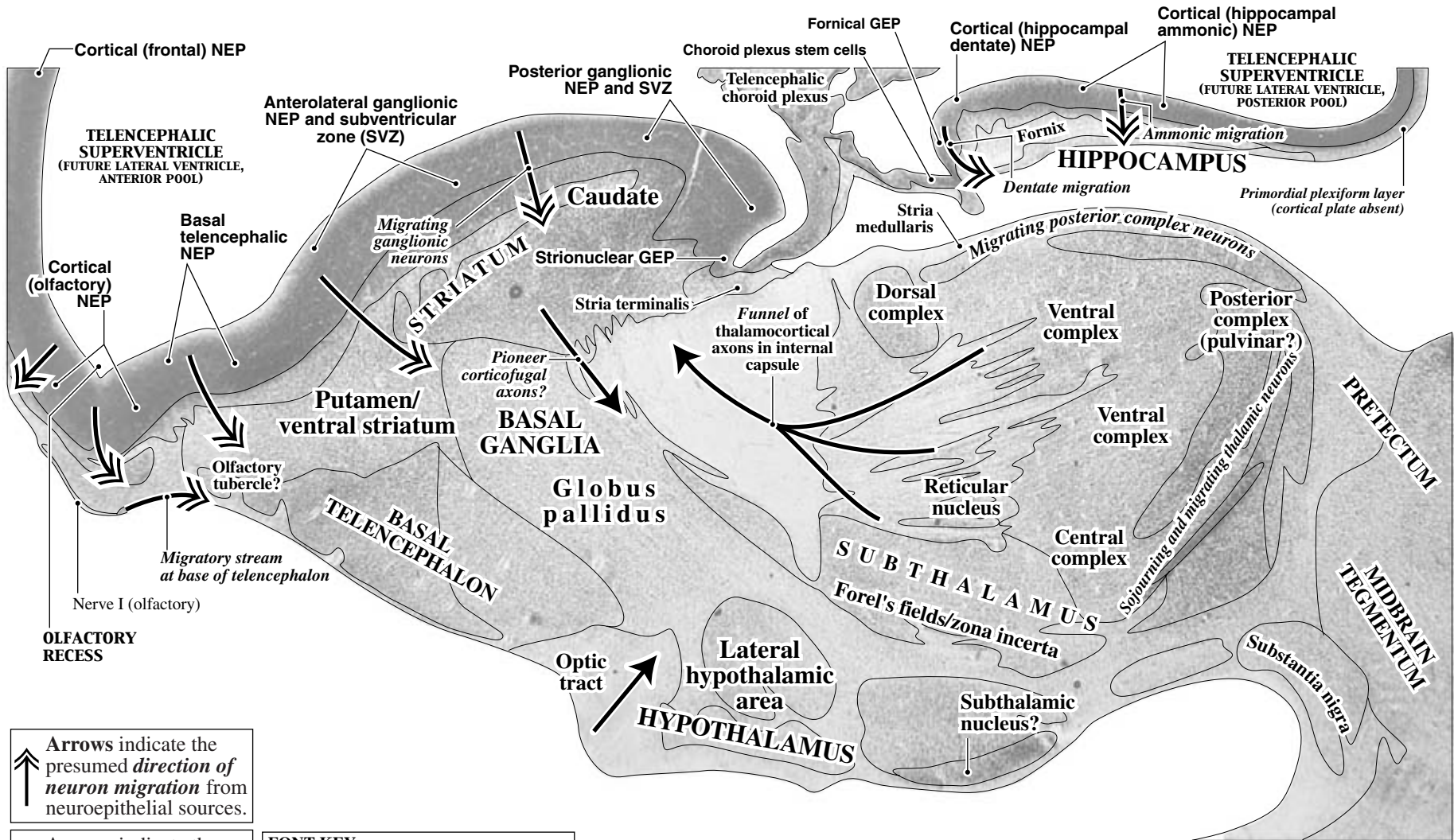
GW8 Sagittal, CR 33 mm, C145
Between levels 5 and 6:
Slide 17, Section 2

BASAL TELENCEPHALON AND DIENCEPHALON



See level 5 in Plates 126A-D, level 6 in Plates 127A-D.

PLATE 137B



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Germinal zone - **Helvetica bold**
 Transient structure - *Times bold italic*
 Permanent structure - **Times Roman or Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

PLATE 138A

**GW8 Sagittal, CR 33 mm, C145
Medial to level 1:
Slide 24, Section 2**

MIDBRAIN TECTUM AND TEGMENTUM



**See level 1 in
Plates 122A-D.**

PLATE 138B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

Dashed lines indicate staining and/or sectioning artifacts.

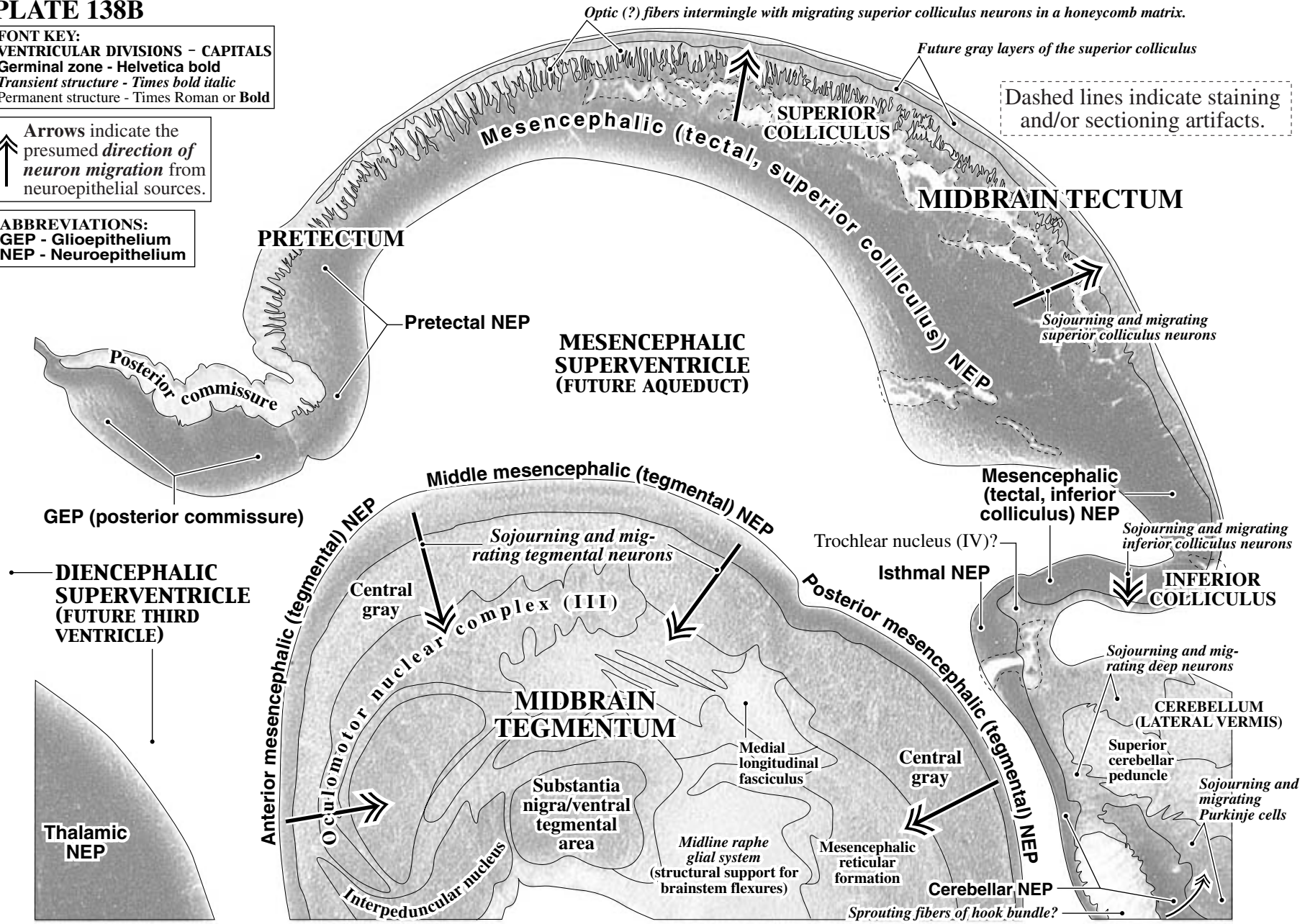
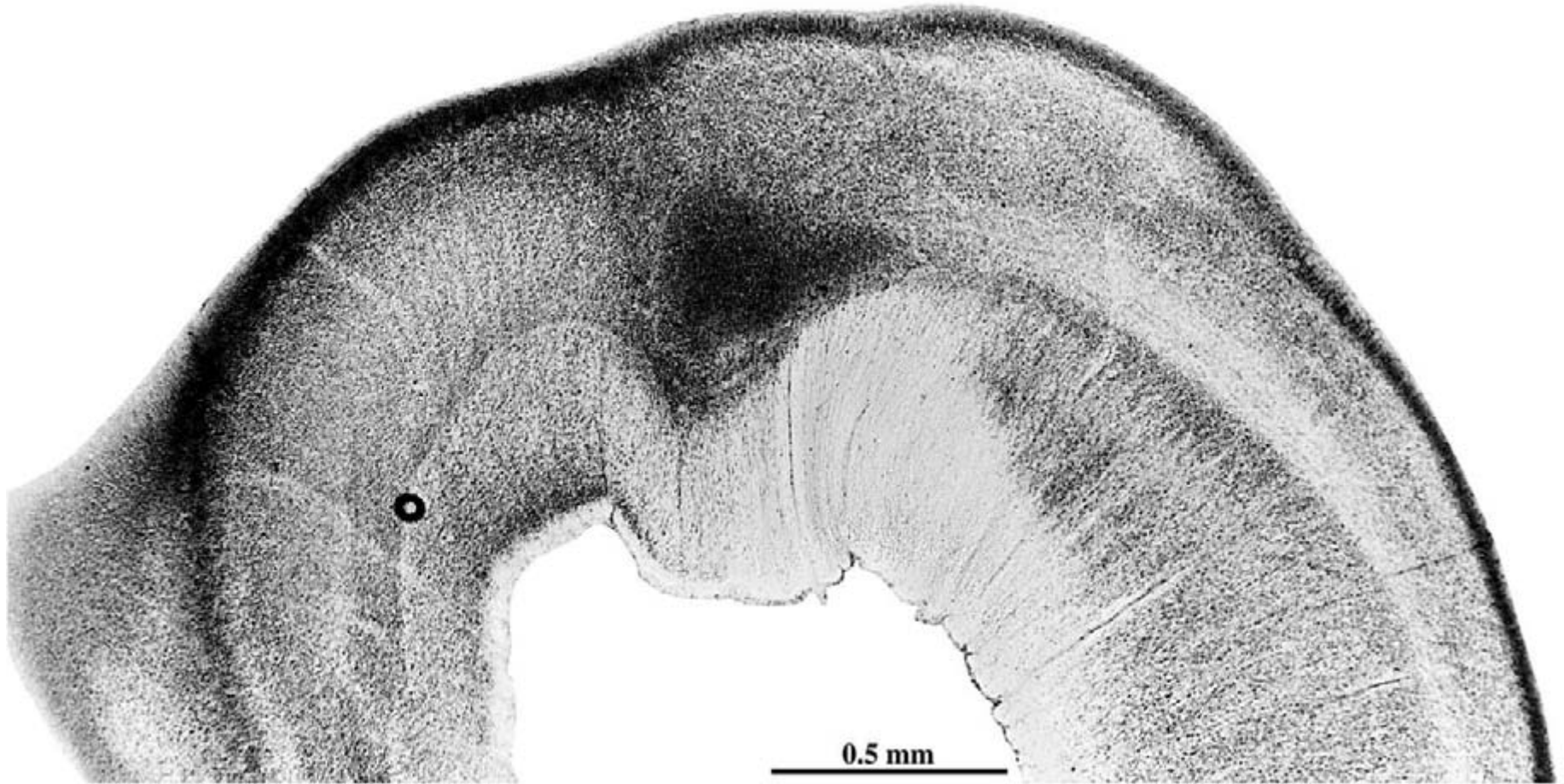


PLATE 139A

**GW8 Sagittal, CR 33 mm, C145
Level 1: Slide 23, Section 2**

MIDBRAIN TEGMENTUM

360



See the entire section in Plates 122A-D.

PLATE 139B

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
*G*erminal zone - Helvetica bold
*T*ransient structure - Times bold *italic*
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

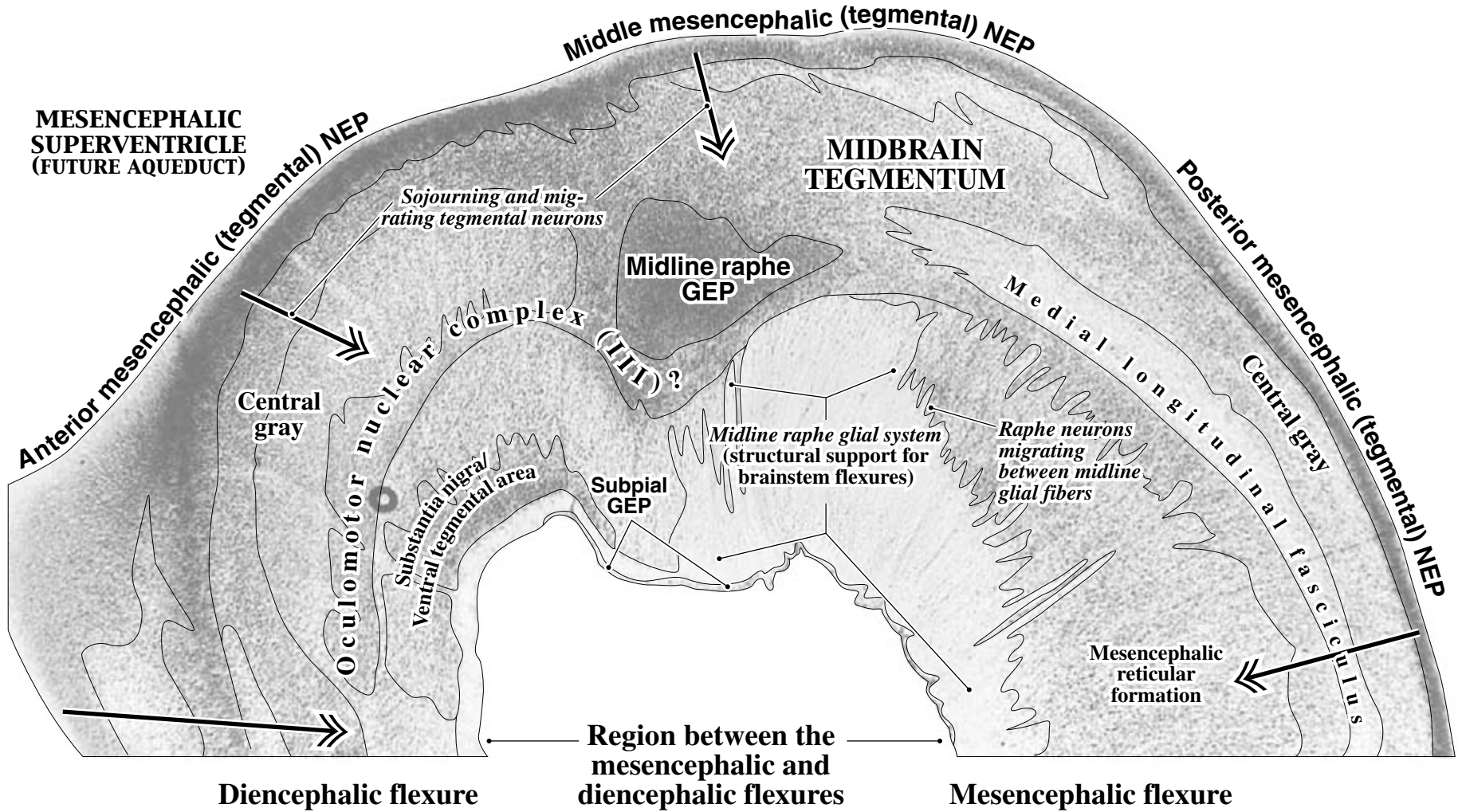
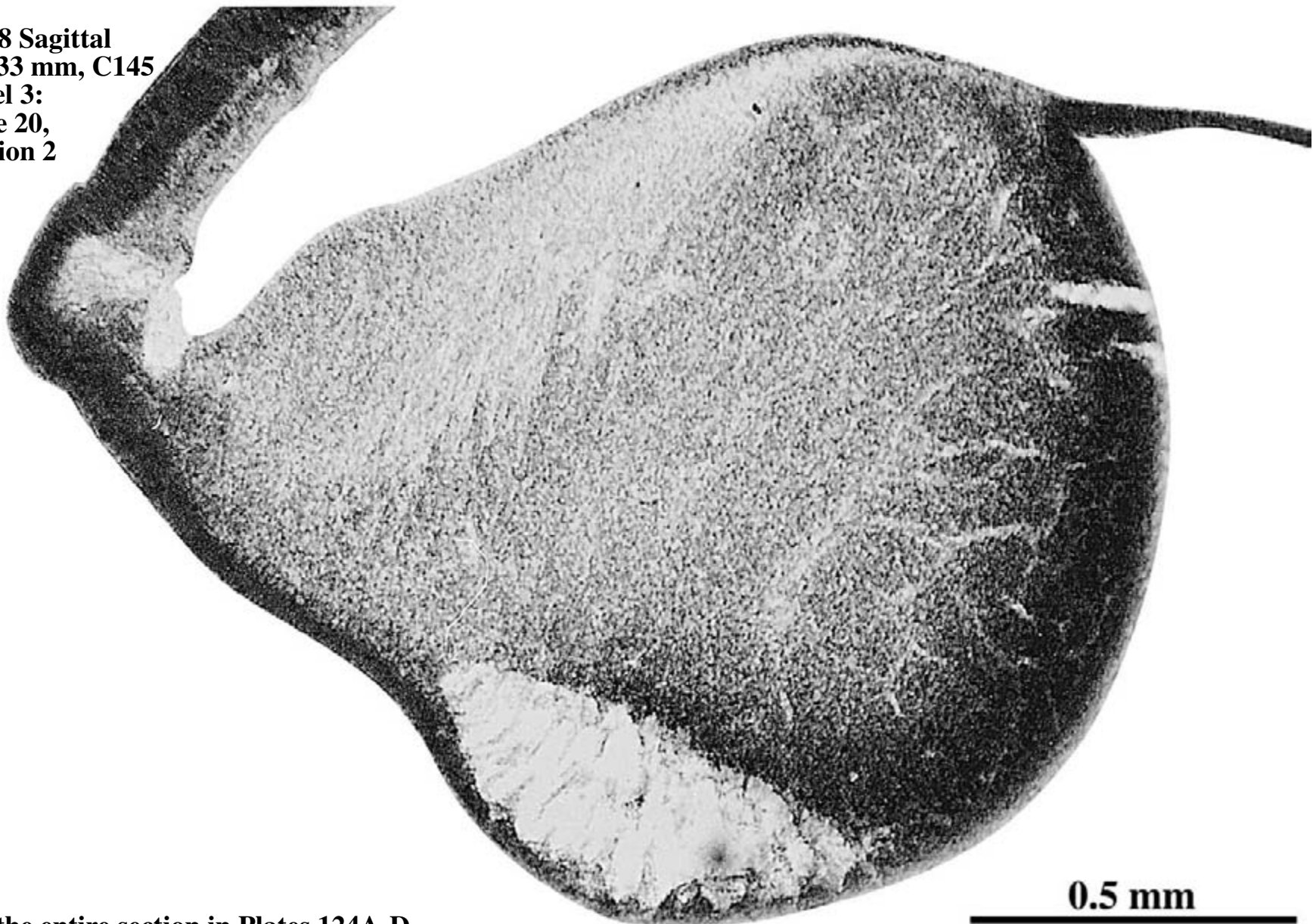


PLATE 140A

CEREBELLUM: LATERAL VERMIS

**GW8 Sagittal
CR 33 mm, C145
Level 3:
Slide 20,
Section 2**



0.5 mm

See the entire section in Plates 124A-D.

PLATE 140B

MESENCEPHALIC SUPERVENTRICLE
(FUTURE AQUEDUCT)

Dashed lines indicate staining
and/or sectioning artifacts.

Dorsal rhombic lip
(contains cerebellar
germinal trigone)

Choroid plexus
stem cells

Medullary
velum

Mesencephalic
(tectal, inferior
colliculus) NEP

INFERIOR
COLLICULUS

External germinal layer

Migrating and
proliferating
external germinal
layer cells

*Sojourning and migrating
inferior colliculus neurons*

Isthmal NEP?

Nerve IV
(trochlear)

*Trajectories of deep
neuron migration*

Superior cerebellar peduncle

CEREBELLUM
(LATERAL VERMIS)

*Trajectory of
Purkinje cell
migration*

*Sojourning and migrating
deep neurons*

Sojourning and migrating deep neurons

Sojourning and migrating Purkinje cells

Cerebellar
(deep neuron) NEP?

Cerebellar
(Purkinje cell)
NEP

*Sprouting fibers of
hook bundle?*

RHOMBENCEPHALIC
SUPERVENTRICLE
(FUTURE FOURTH VENTRICLE)

NEP - Neuroepithelium

Arrows indicate the
presumed *direction of
neuron migration* from
neuroepithelial sources.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 141A

CEREBELLUM: HEMISPHERE

**GW8 Sagittal
CR 33 mm, C145
Level 9: Slide 12, Section 4**



See the entire section in Plates 130A-D.

PLATE 141B

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

NEP - Neuroepithelium

Dashed lines indicate staining and/or sectioning artifacts.

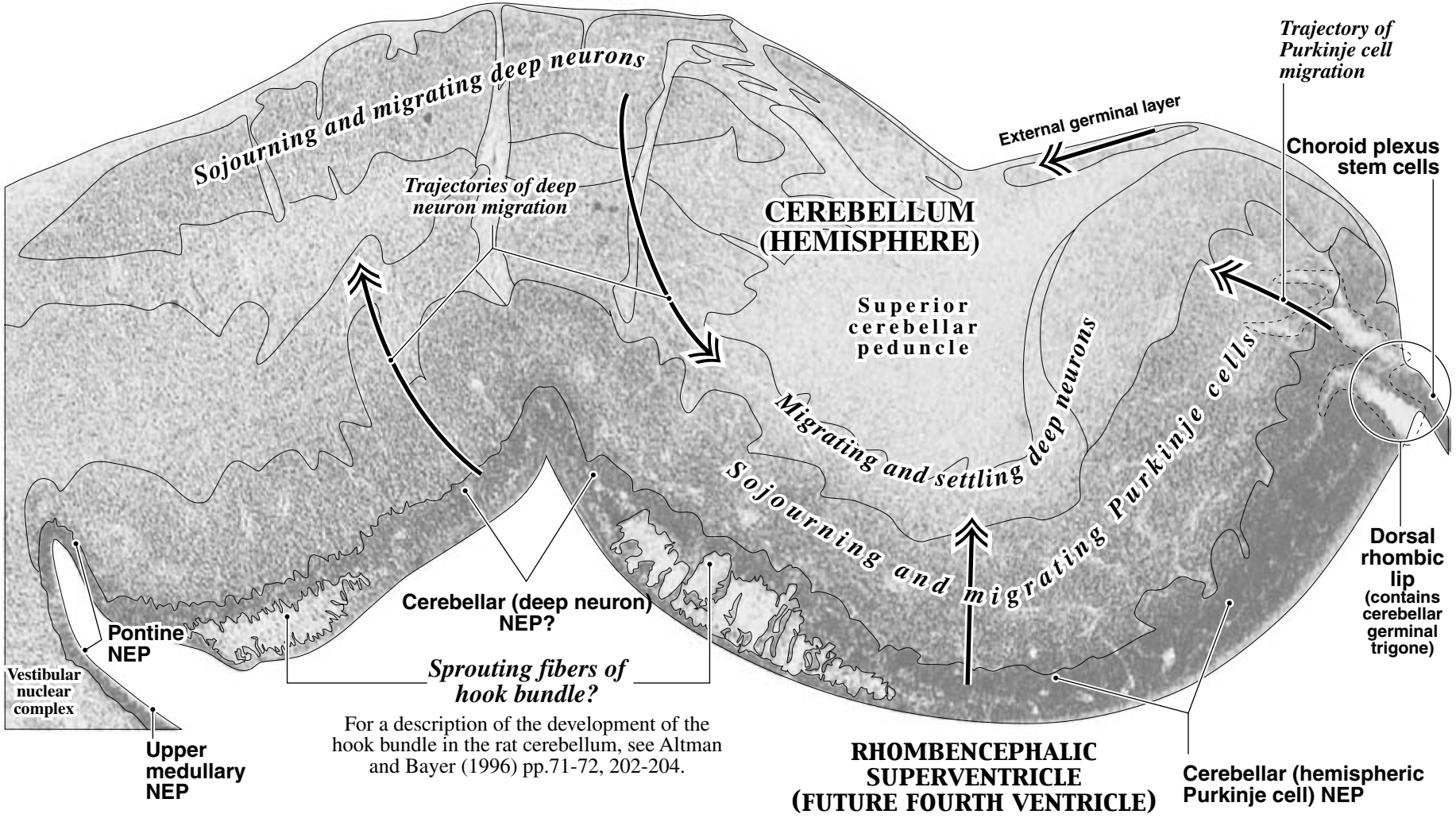
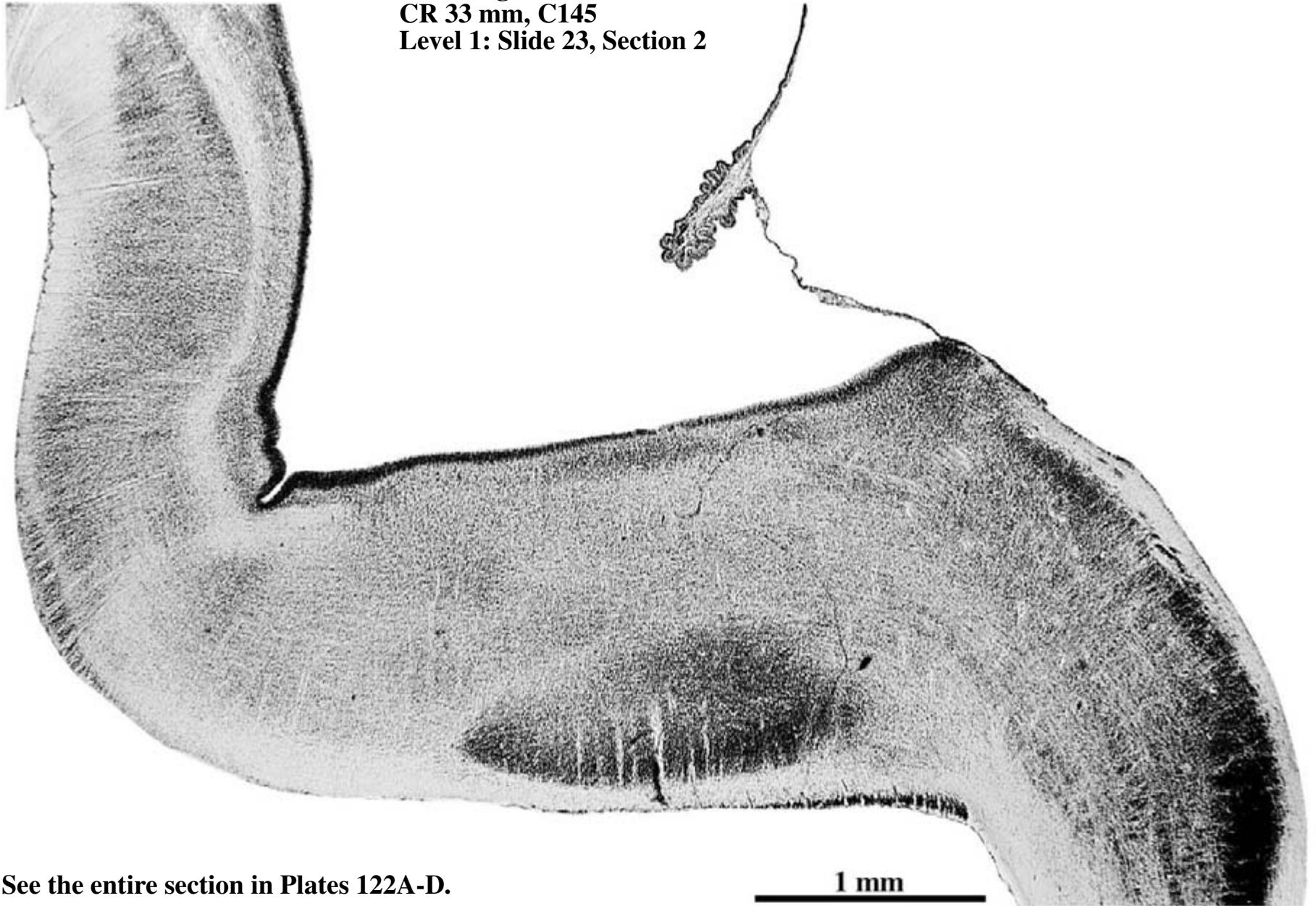


PLATE 142A

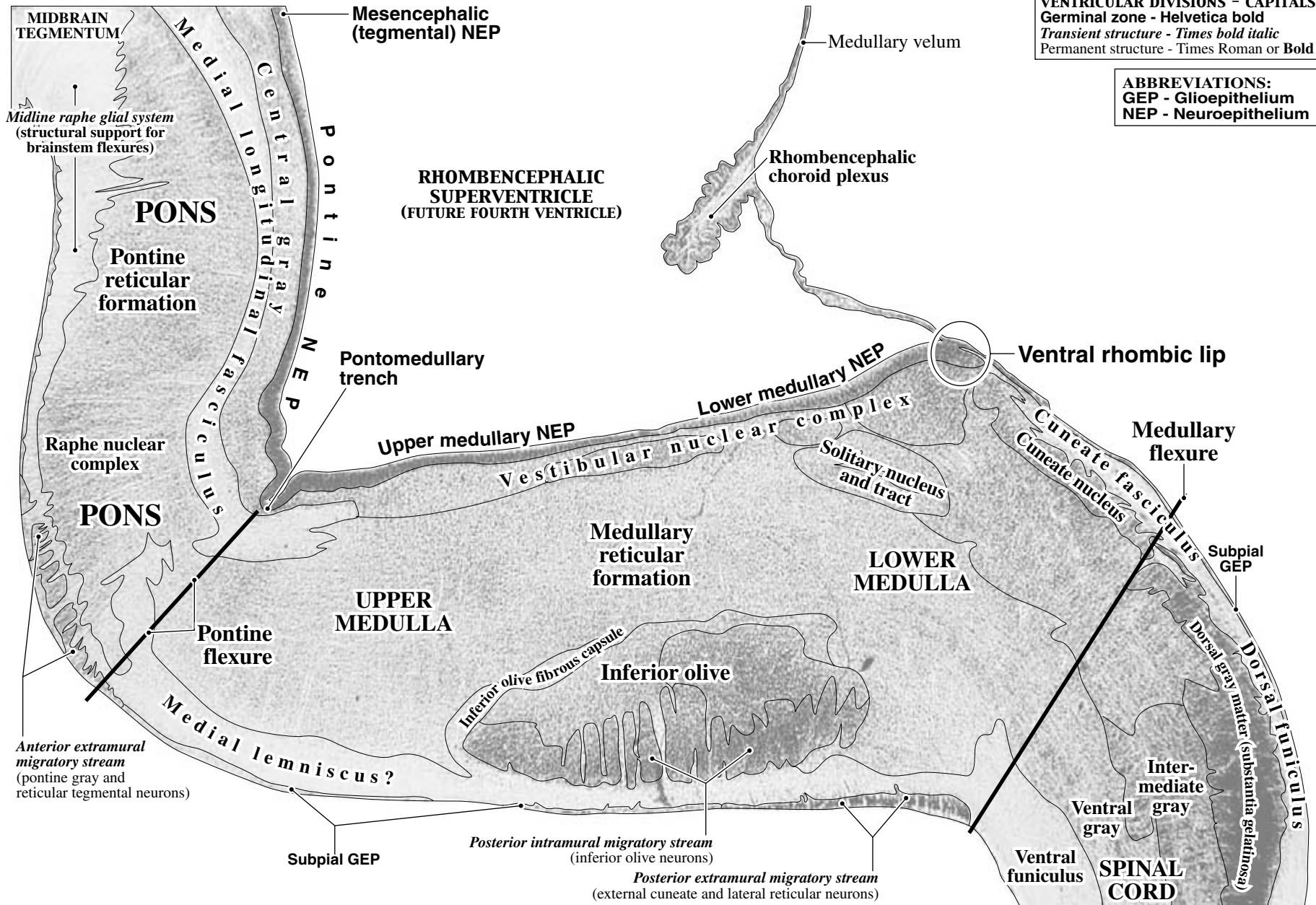
**GW8 Sagittal
CR 33 mm, C145
Level 1: Slide 23, Section 2**

PONS AND MEDULLA



See the entire section in Plates 122A-D.

PLATE 142B



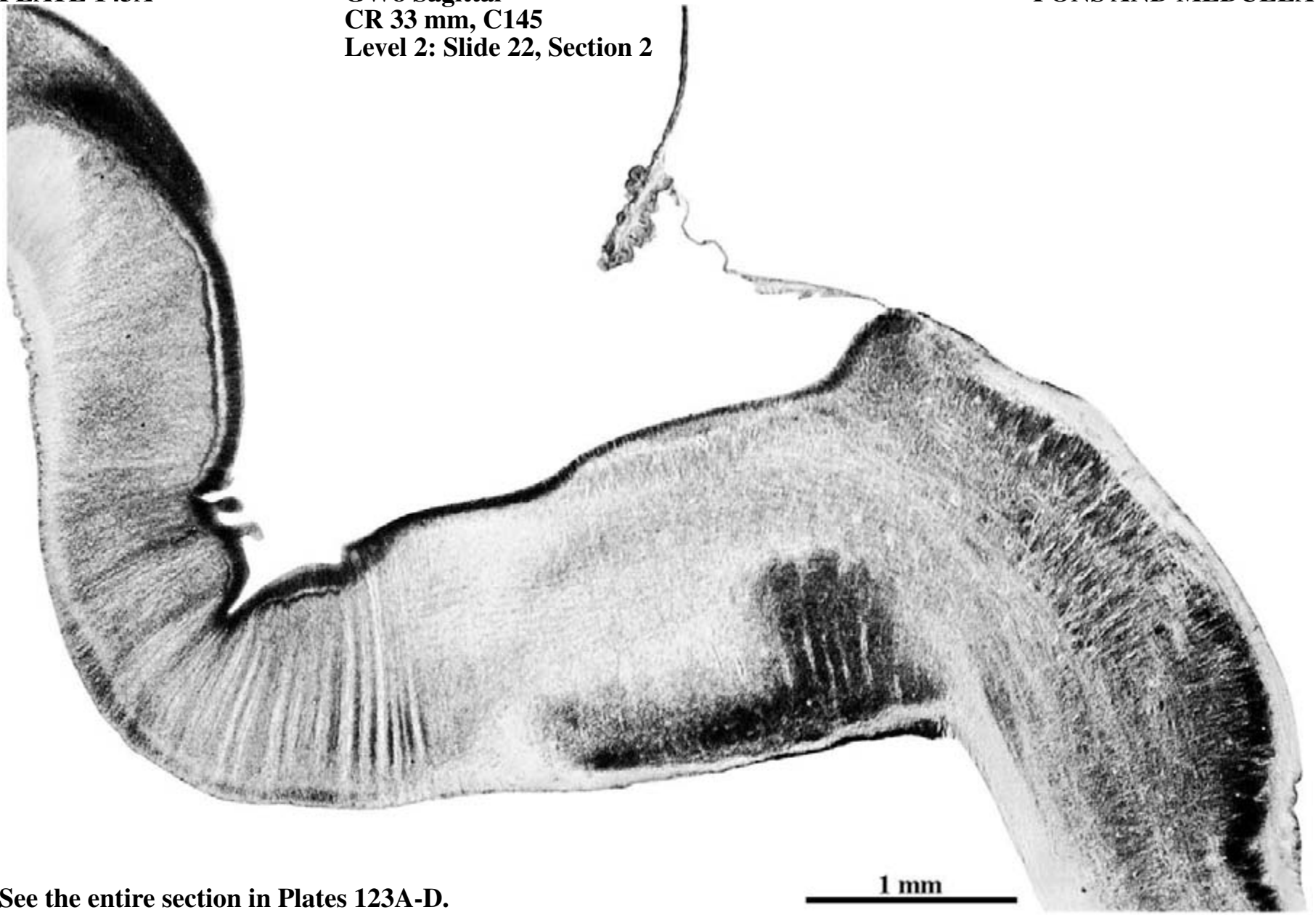
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

PLATE 143A

**GW8 Sagittal
CR 33 mm, C145
Level 2: Slide 22, Section 2**

PONS AND MEDULLA



See the entire section in Plates 123A-D.

PLATE 143B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

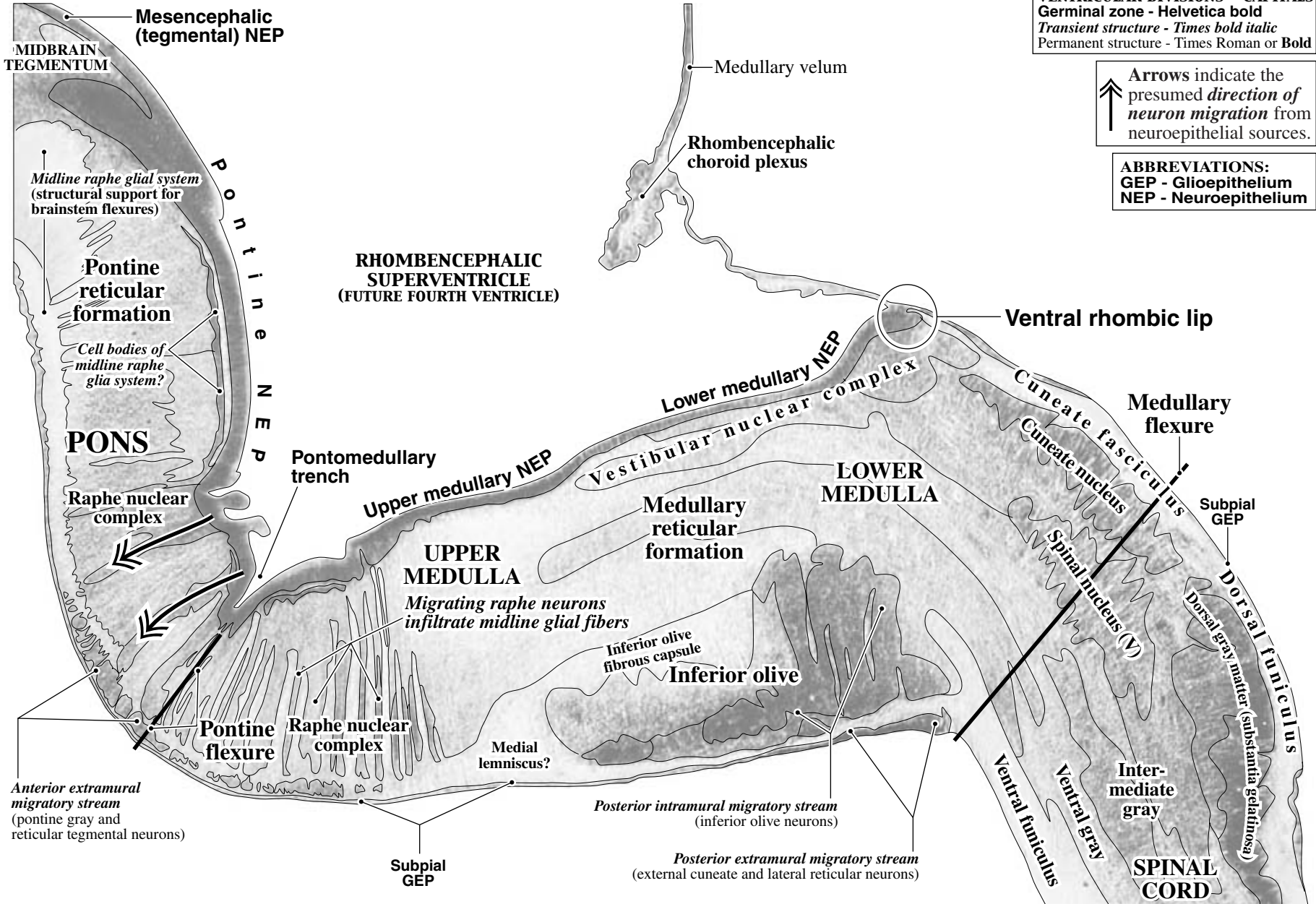
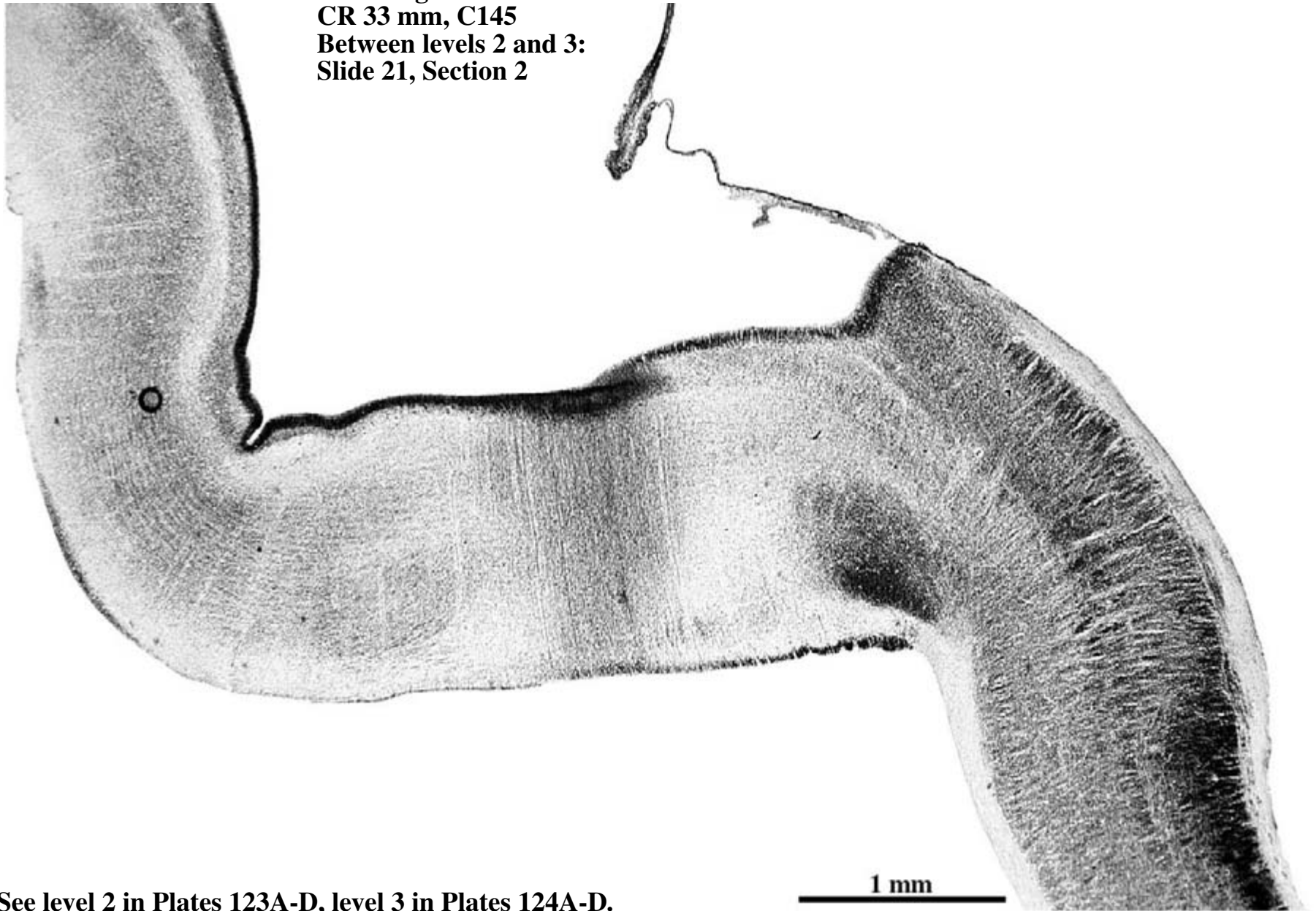


PLATE 144A

**GW8 Sagittal
CR 33 mm, C145
Between levels 2 and 3:
Slide 21, Section 2**

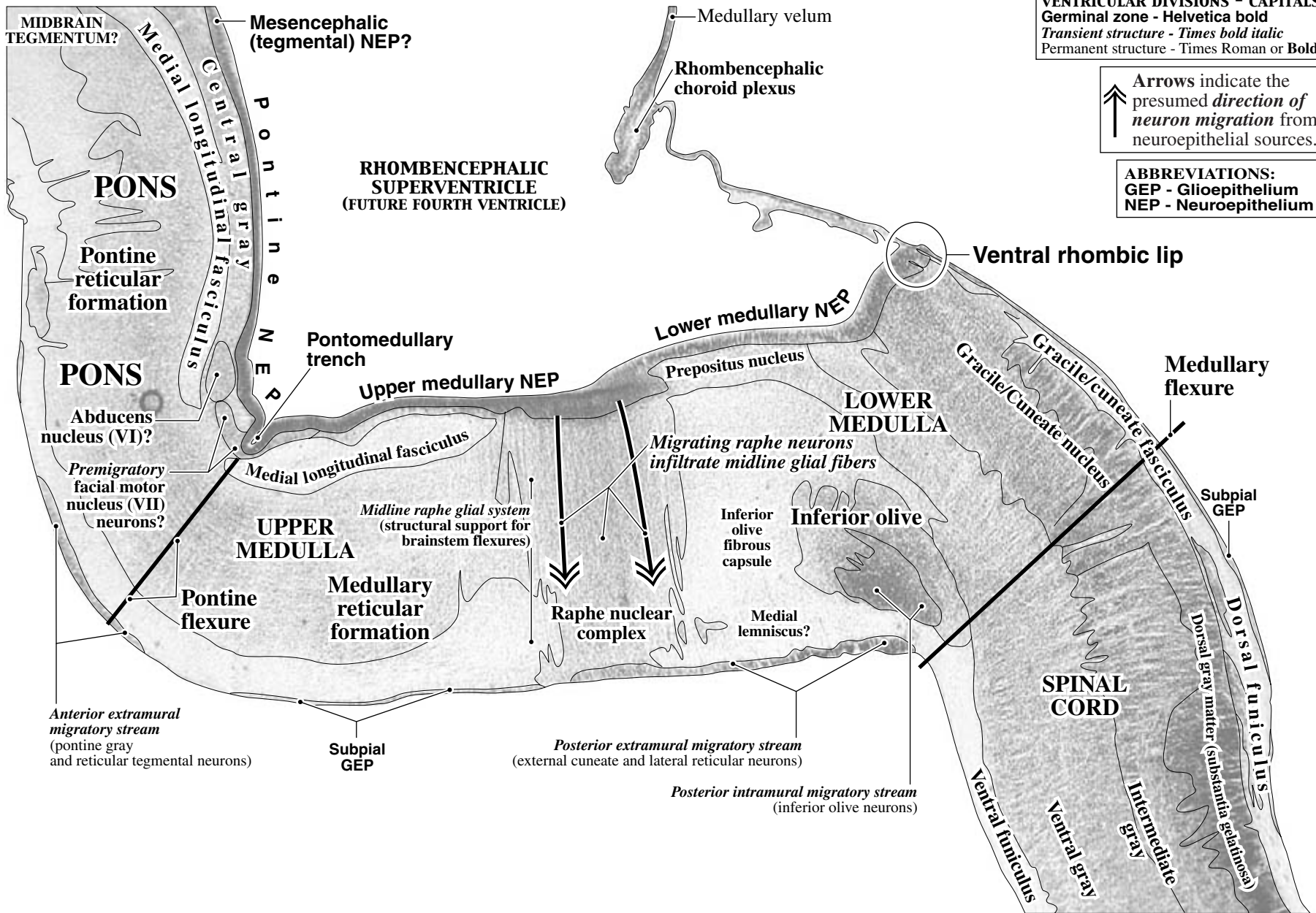
BRAINSTEM



See level 2 in Plates 123A-D, level 3 in Plates 124A-D.

1 mm

PLATE 144B



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

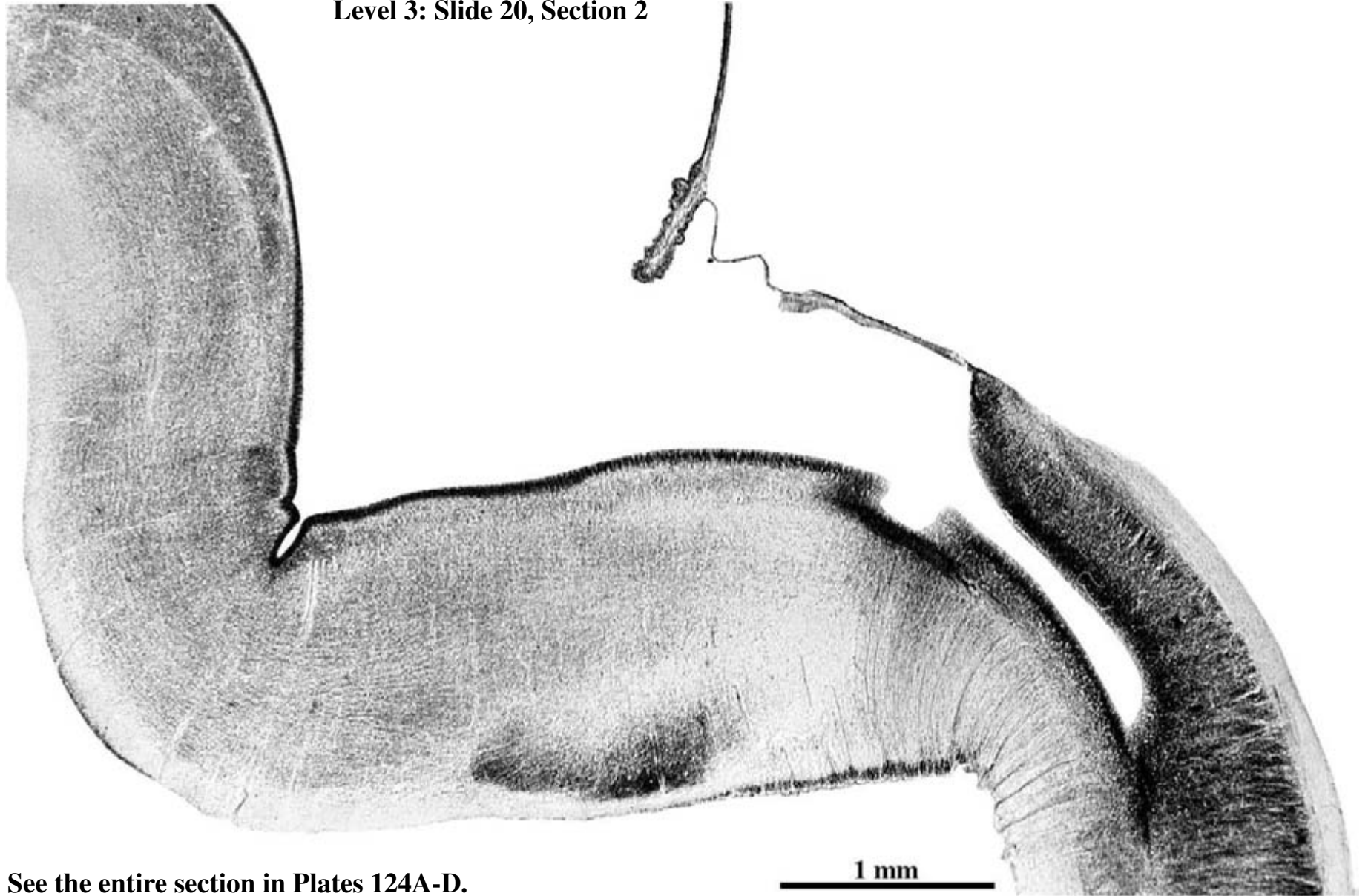
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

PLATE 145A

**GW8 Sagittal
CR 33 mm, C145
Level 3: Slide 20, Section 2**

BRAINSTEM



See the entire section in Plates 124A-D.

PLATE 145B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

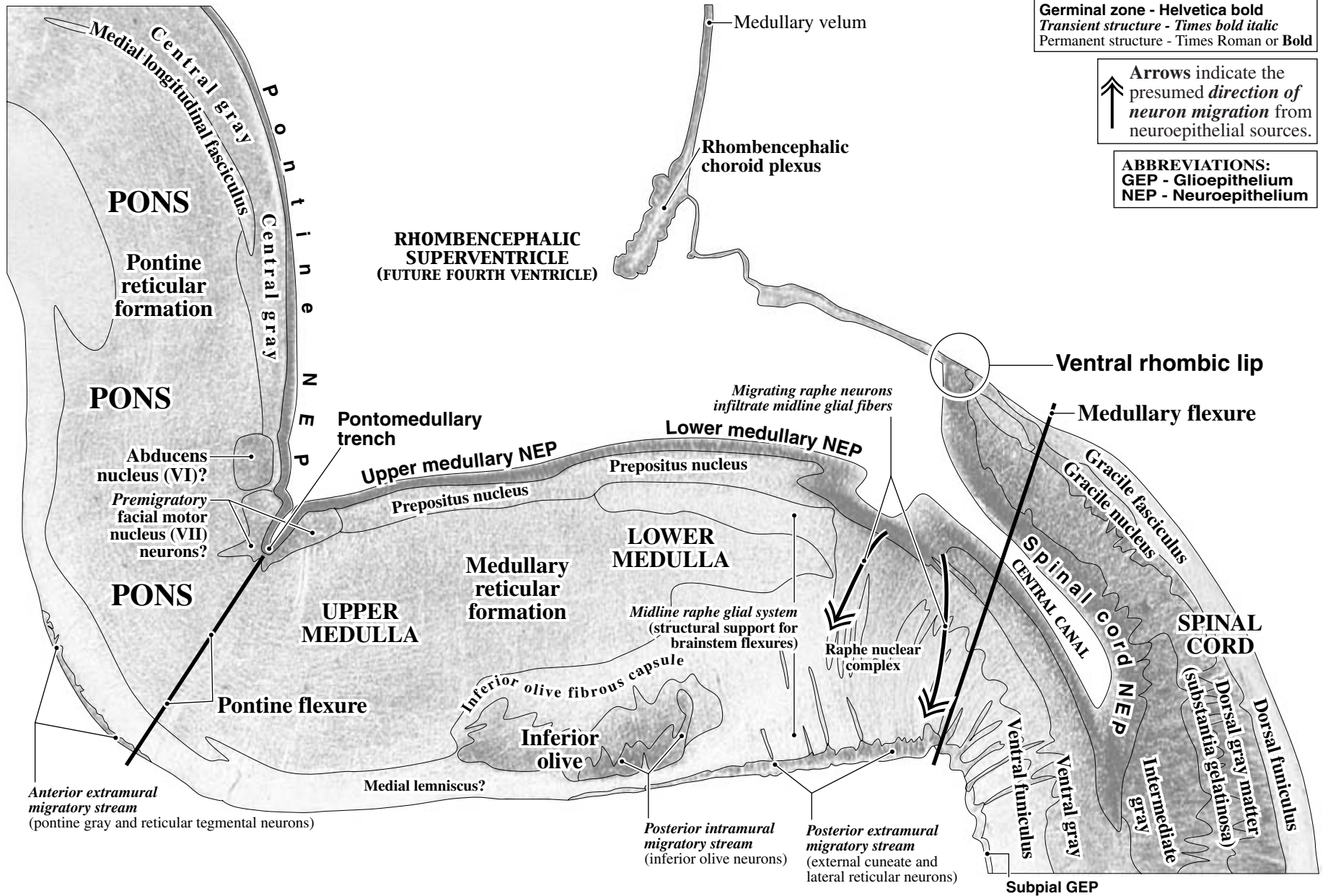
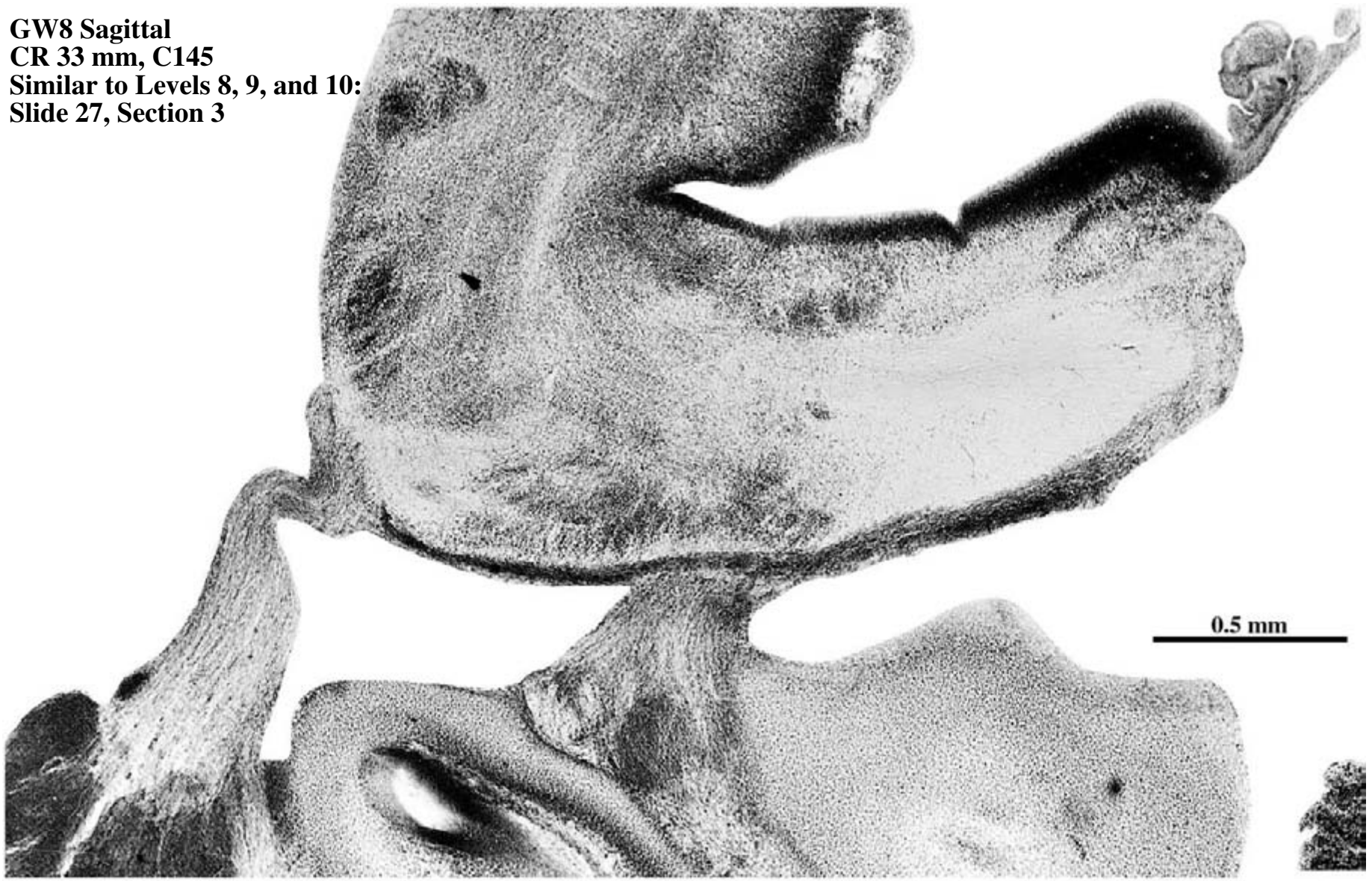


PLATE 146A

LATERAL PONS, MEDULLA, AND SENSORY GANGLIA

**GW8 Sagittal
CR 33 mm, C145
Similar to Levels 8, 9, and 10:
Slide 27, Section 3**



See similar areas from the left side of the brain in Plates 129A-D to 132A-D.

PLATE 146B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

Neuroepithelium - NEP

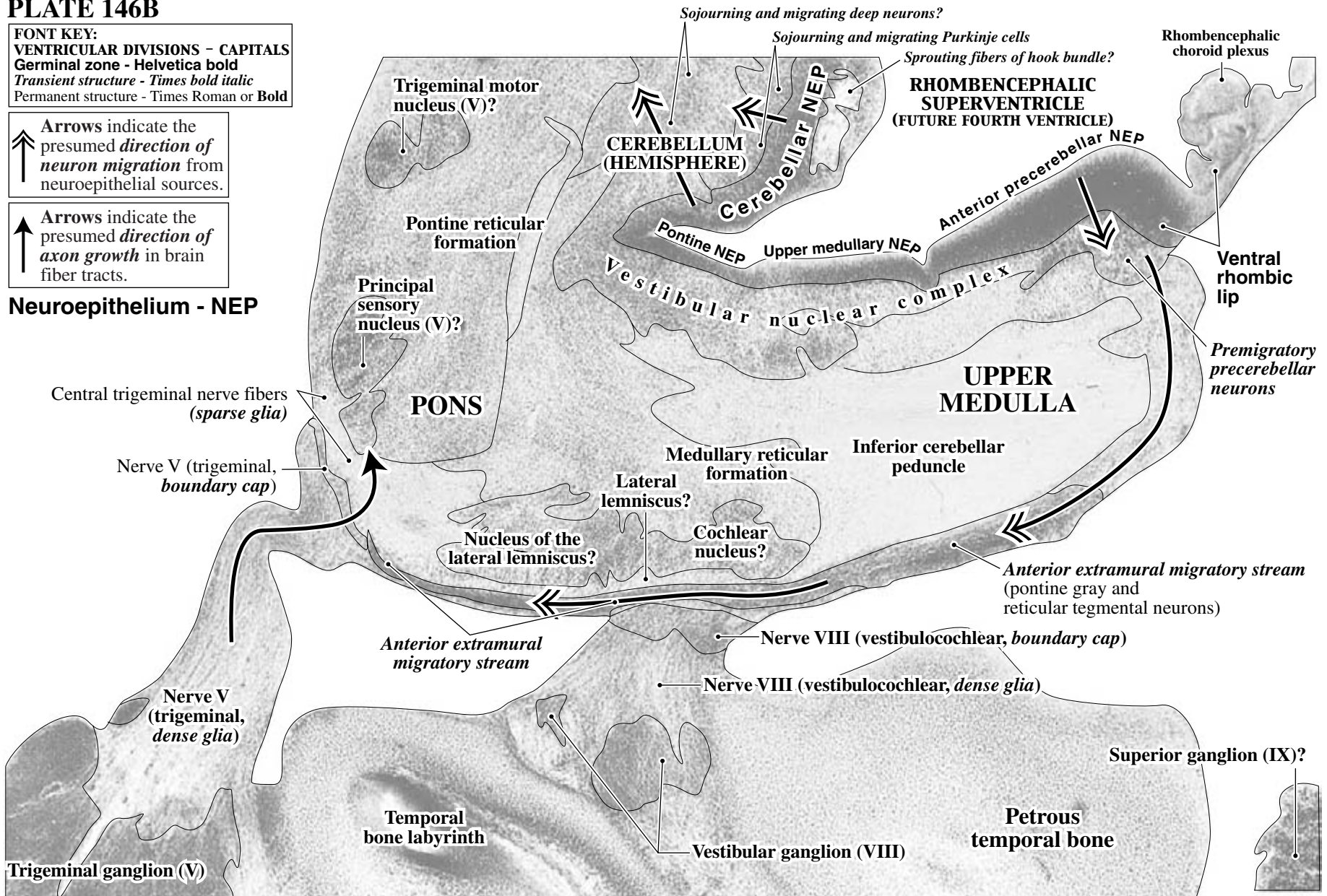
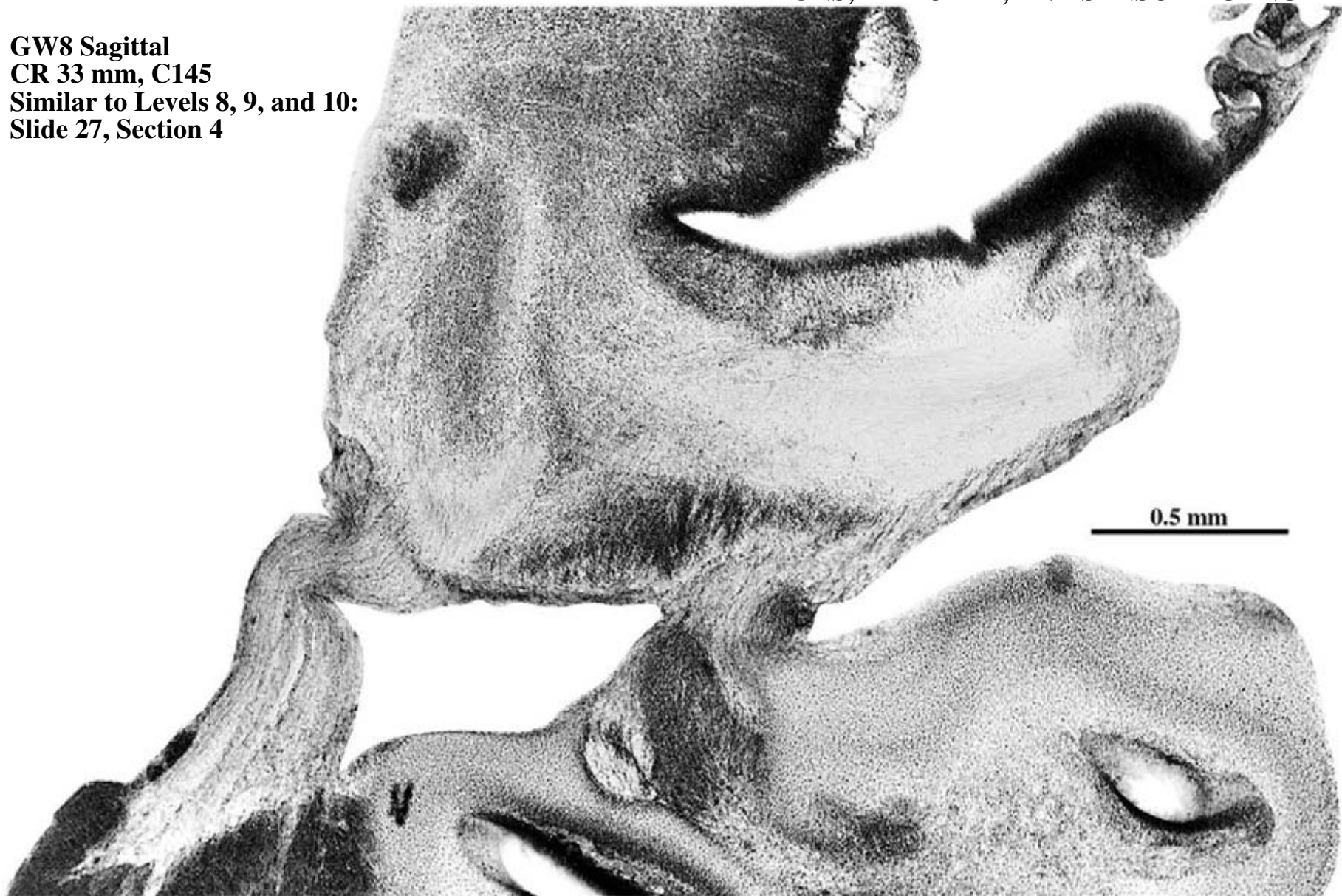


PLATE 147A

LATERAL PONS, MEDULLA, AND SENSORY GANGLIA

**GW8 Sagittal
CR 33 mm, C145
Similar to Levels 8, 9, and 10:
Slide 27, Section 4**



See similar areas from the left side of the brain in Plates 129A-D to 132A-D.

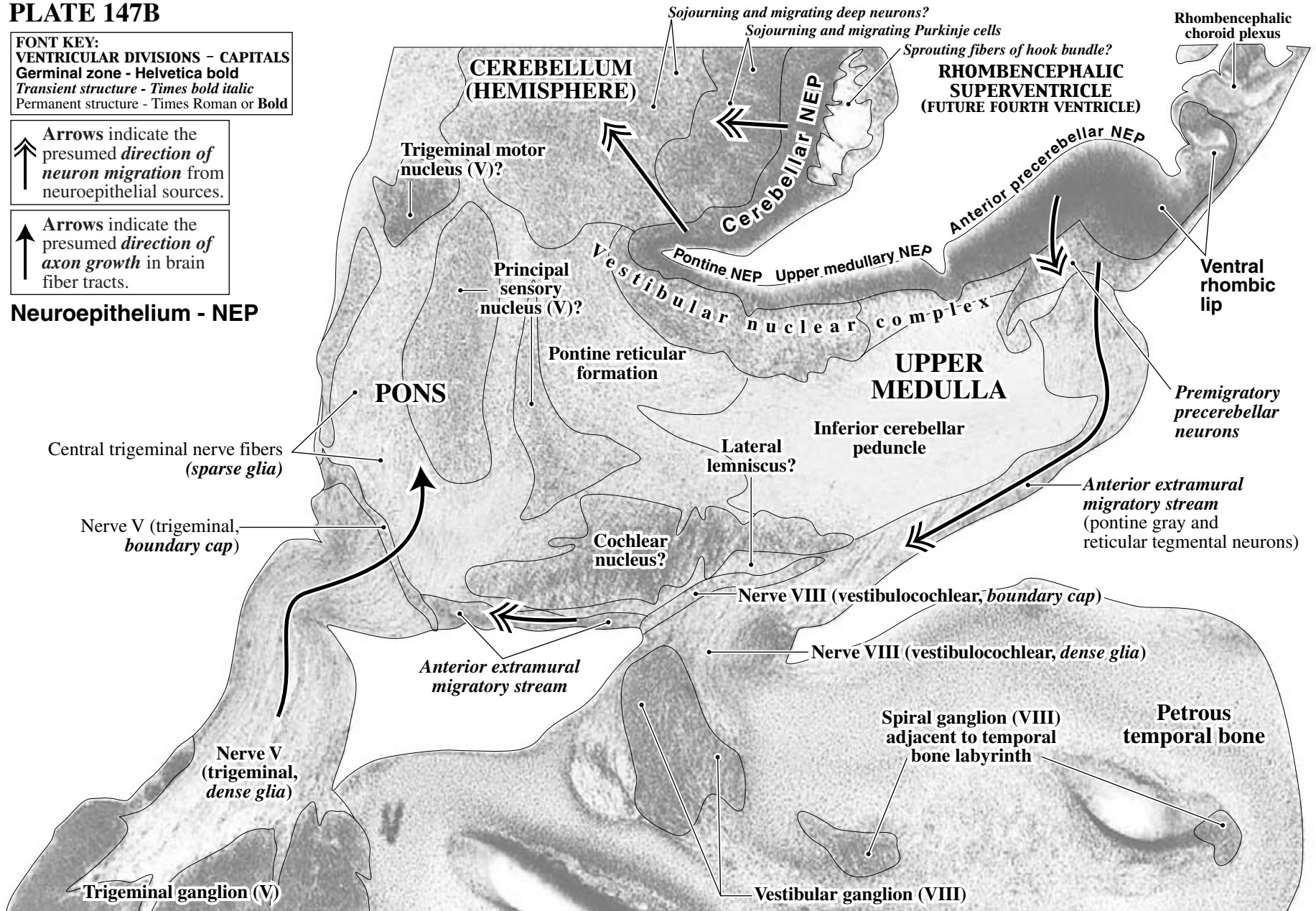
PLATE 147B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

Neuroepithelium - NEP



PART VIII: GW8 CORONAL

This is specimen number 9226 in the Carnegie Collection, designated here as C9226. A normal male fetus with a crown-rump length (CR) of 31 mm was collected in 1954, and is estimated to be in gestational week (GW) 8. The entire fetus was embedded in paraffin mixed with 8% celloidin, cut transversely in 10 μ m thick sections, and stained with azan. The histology of this specimen is remarkable, and the sections are nearly perfectly bilateral. Since there is no photograph of this brain before it was embedded and cut, a specimen from Hochstetter (1919) that is only partially comparable to C9226 has been modified to show the approximate section plane and external features of the brain at GW8 (**Figure 7**). Like most of the specimens in this Volume, the sections are not cut exactly in one plane; C9226's cortex is cut midway between coronal and horizontal planes. Since the cerebral cortex is in every section and the brainstem is cut in a more horizontal orientation, the brain more closely resembles a coronally sectioned brain. Unfortunately, the Hochstetter specimen is less mature (CR27 mm) and we could not find a drawing of a brain specimen that would fit C9226. The C9226 sections through the cortex and brainstem are not in the same plane when transferred to Hochstetter's CR27 mm specimen. Instead, brainstem planes of section appear to fan upward and downward from sections in the cortex. We interpret this to indicate that the brain flexures are more loosely folded in the Hochstetter specimen than in C9226. If one "squeezes" the brainstem to make the folds tighter, the cortex and brainstem planes would line up. Photographs of 23 sections (**Levels 1-23**) are illustrated at low magnification in **Plates 148-167**. High-magnification views of different areas of the cerebral cortex are shown in **Plates 168-169**.

C9226 is similar to the other GW8 specimens and shows brain maturation in a different perspective. Each of the brain's major subdivisions has a large *supraventricle* in the cores, especially the telencephalon and the rhombencephalon. Midline sagittal sections have large diencephalic and mesencephalic supraventricles because the cuts are parallel to their dorsoventral and anteroposterior axes (see C145 in **Part VII**). C9226's coronal sections show the slit-like shapes of the diencephalic and mesencephalic supraventricles in the midline.

The parenchyma, the area between the superficial border of the *neuroepithelium* (NEP) / *subventricular zone* (SVZ) and the pial membrane, is the region where neurons migrate, settle, and differentiate. The thicknesses of the neuroepithelium and the parenchyma are clues to the level of maturation of a developing brain structure.

The parenchyma is thick and bordered by a thin NEP in the medulla, pons, and midbrain tegmentum, indicating that most neurons have been generated in these structures. Furthermore, the lack of dense accumulations of cells just outside the NEP in the midbrain tegmentum, pons, and medulla indicate that very few neurons are being generated, few are migrating, and most are settled and differentiating. There are two exceptions in the medulla and pons. First, near the pontomedullary trench, presumptive facial motor neurons are migrating toward their ventral pontine/medullary settling sites. Second, the *precerebellar neuroepithelium* in the medulla is thicker and generating pontine gray (and possibly other neurons); many precerebellar neurons are migrating in the *anterior and posterior extramural migratory streams*. The cerebellar NEP is thicker than that in the pons and medulla. The cerebellar parenchyma contains a very dense Purkinje cell sojourn zone outside the NEP and presumptive earlier-generated deep neurons lie in a superficial position. Like C145, the *external germinal layer* (*egl*) is barely visible emanating from the germinal trigone in the dorsal rhombic lip. The mesencephalic tectal NEP is thicker than the tegmental NEP and its very thin parenchyma contains dense sojourning and migrating tectal neurons adjacent to the NEP. The tectum is one of the most immature brain structures.

The diencephalic NEP is thicker indicating that many neurons are still being generated even though there is also a thick parenchyma, especially in the thalamus. That is because the thalamus is very large in the mature human brain. There are dense accumulations of young neurons in sojourn zones outside the hypothalamic and thalamic NEPs, indicating that cell migration is more active than final settling and differentiation.

Within the telencephalon, the cerebral cortex has a thick NEP and a very thin parenchyma, indicating that it is the most immature brain structure. The cerebral cortical NEP is the sole germinal matrix. The *stratified transitional field* (*STF*) contains *STF1* and *STF5* only in lateral areas. The pronounced anterolateral (thicker) to dorsomedial (thinner) maturation gradient is evident in both the *cortical plate* and the *STF* layers. In contrast, both the basal telencephalic NEP/SVZ and parenchyma are thick. That is because the basal telencephalon contains many early-generated neuronal populations (for example, globus pallidus and substantia innominata) and massive late-generated populations (striatal neurons in the caudate and putamen). Most of the neurons settling in the basal telencephalon at GW8 are those of the early-generated populations.

GW8 "CORONAL" SECTION PLANES

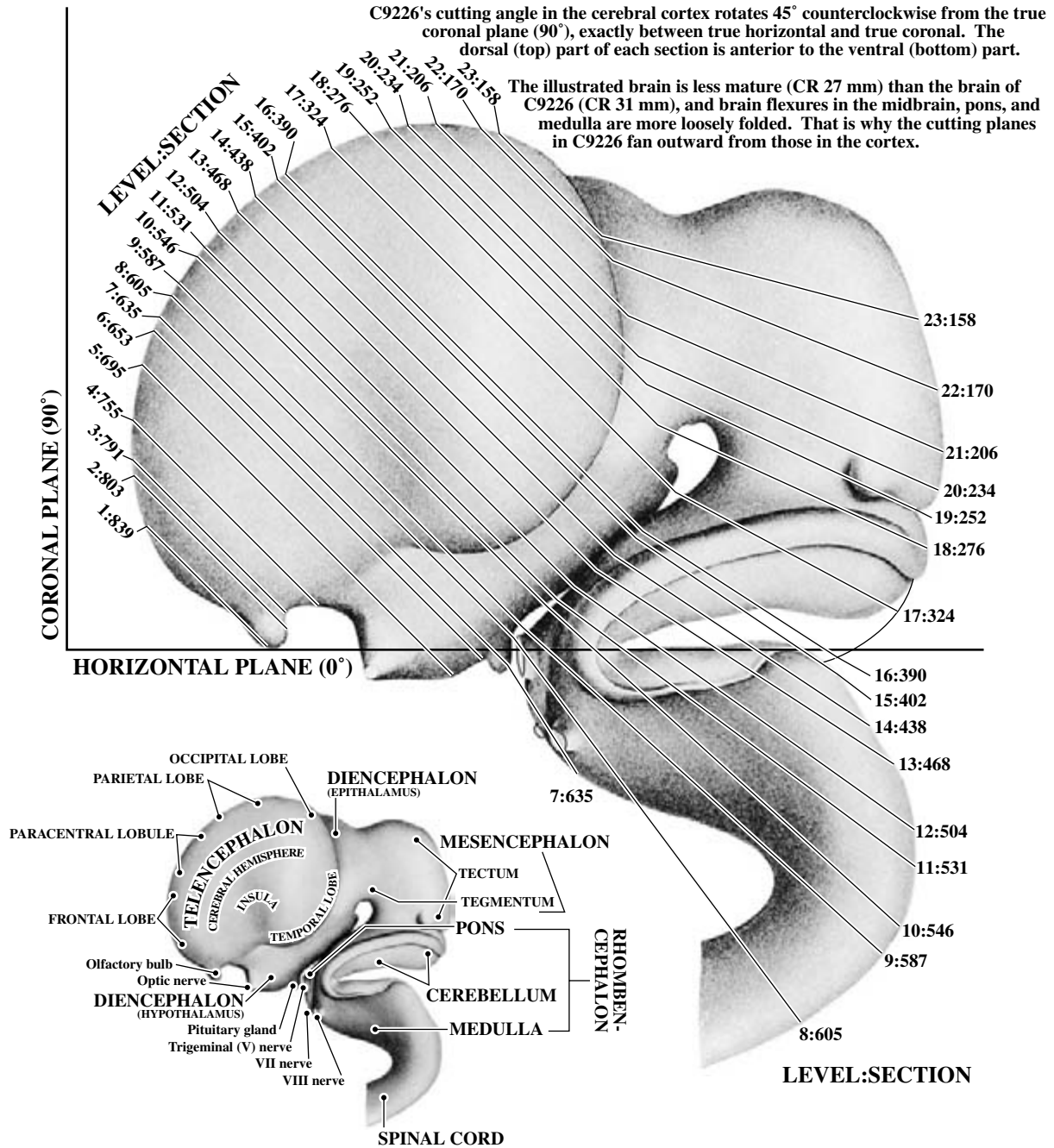
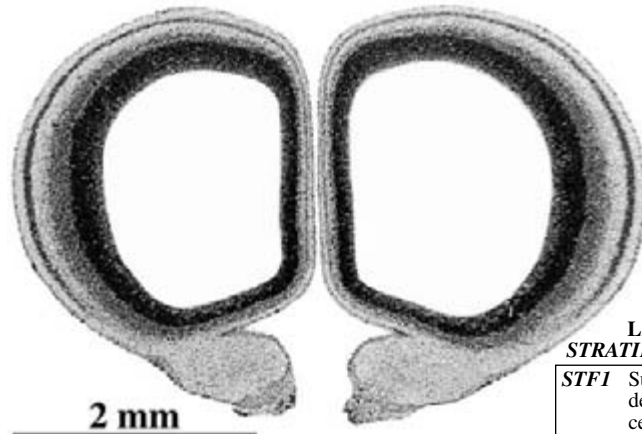


Figure 7. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 27 mm (modified from Figure 37, Table VII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of C9226 in the following pages. The small inset identifies the major structural features. The line in the cerebellum and dorsal edges of the pons and medulla is the cut edge of the medullary velum.

PLATE 148A

GW8 Coronal
CR 31 mm
C9226

Level 1: Section 820

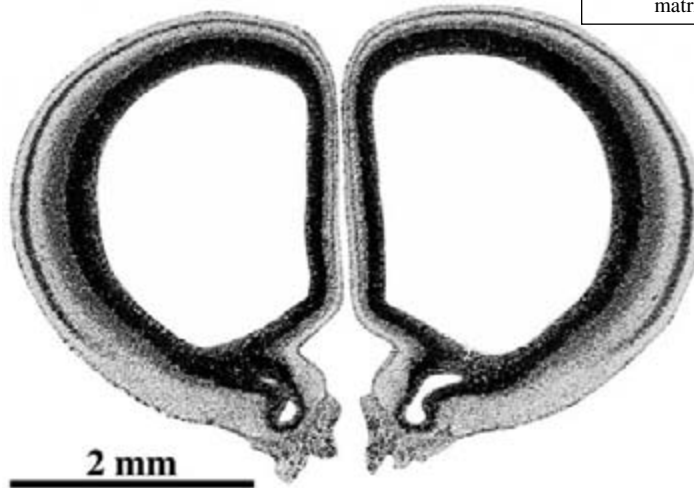


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

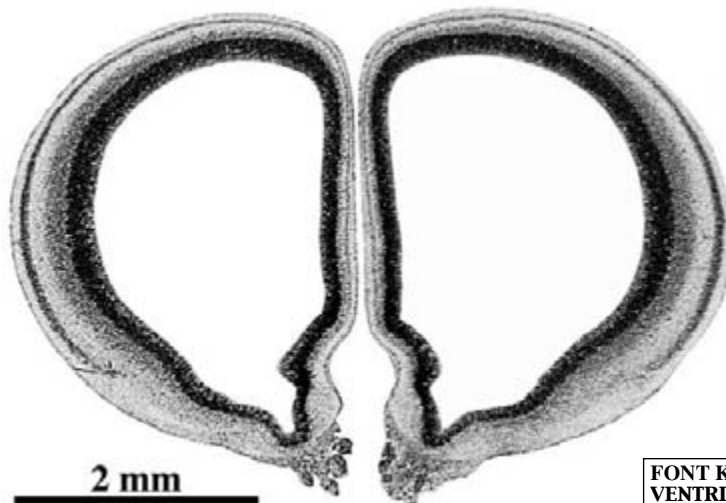
STF1 Superficial fibrous layer with an early developmental stage (*I1*) when many cells are migrating through it, followed by a late stage (*I2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

Level 2: Section 803

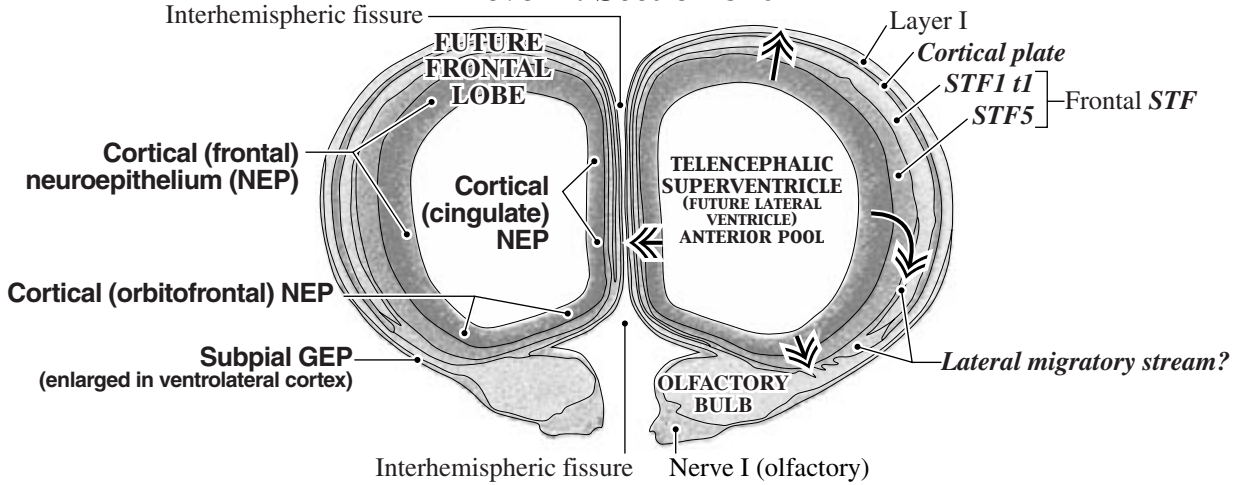


Level 3: Section 791

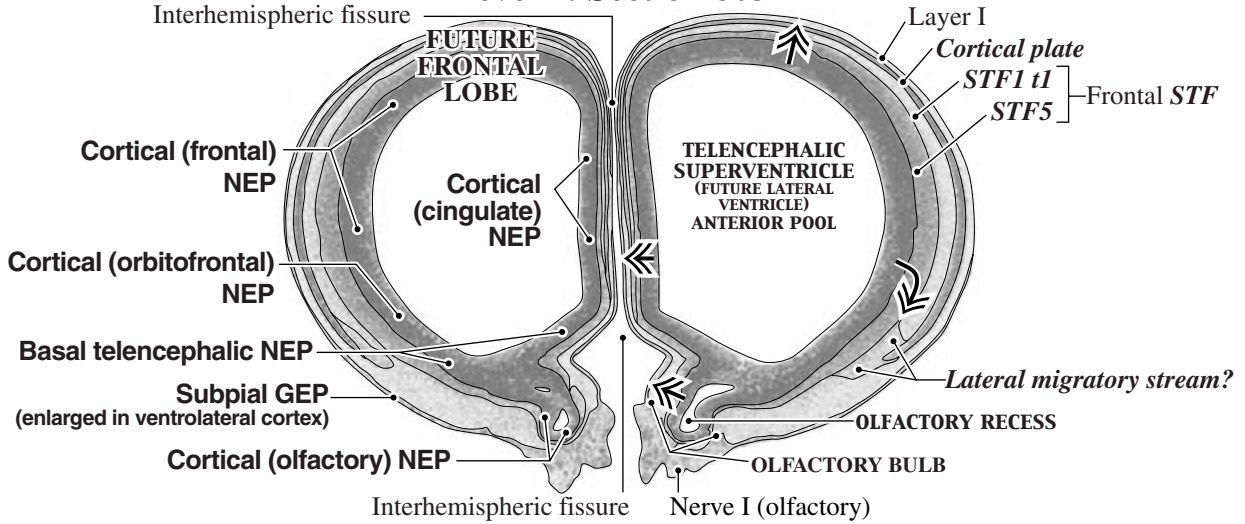


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

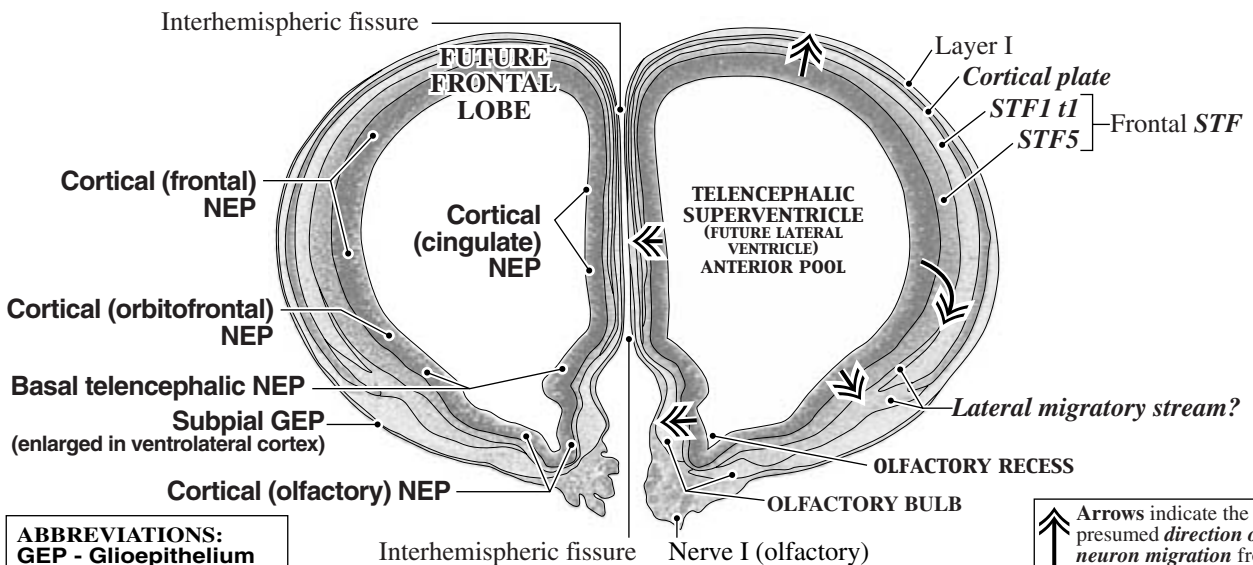
Level 1: Section 820



Level 2: Section 803



Level 3: Section 791



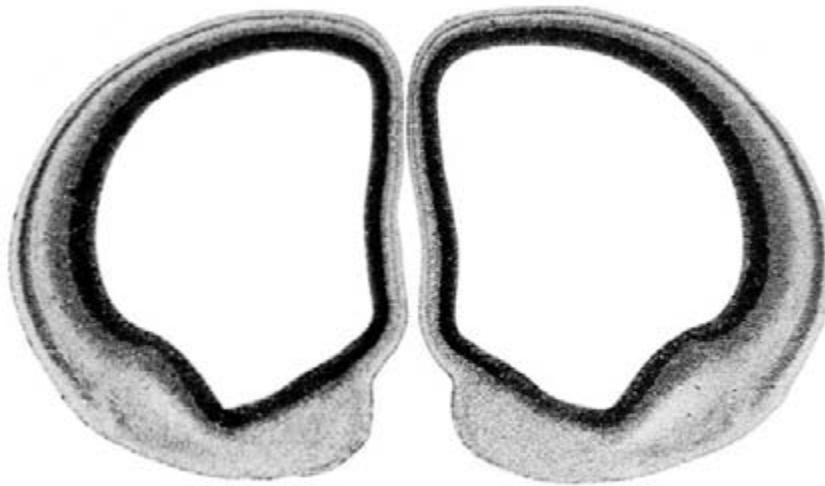
ABBREVIATIONS:
 GEP - Glioepithelium
 NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 149A

GW8 Coronal
CR 31 mm
C9226

Level 4: Section 755



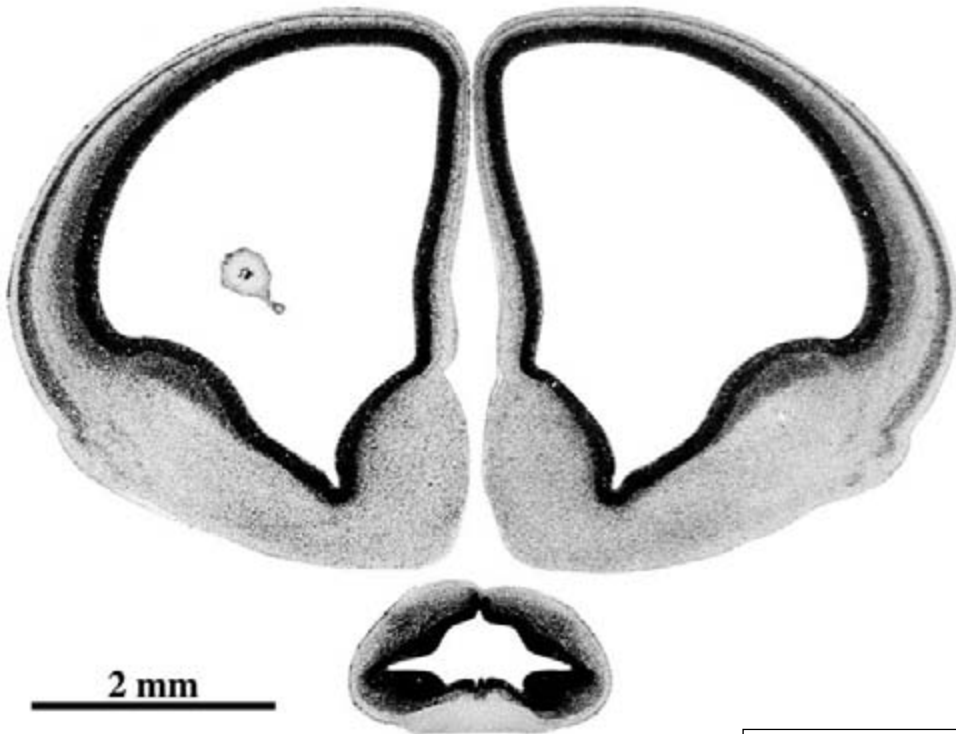
2 mm

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

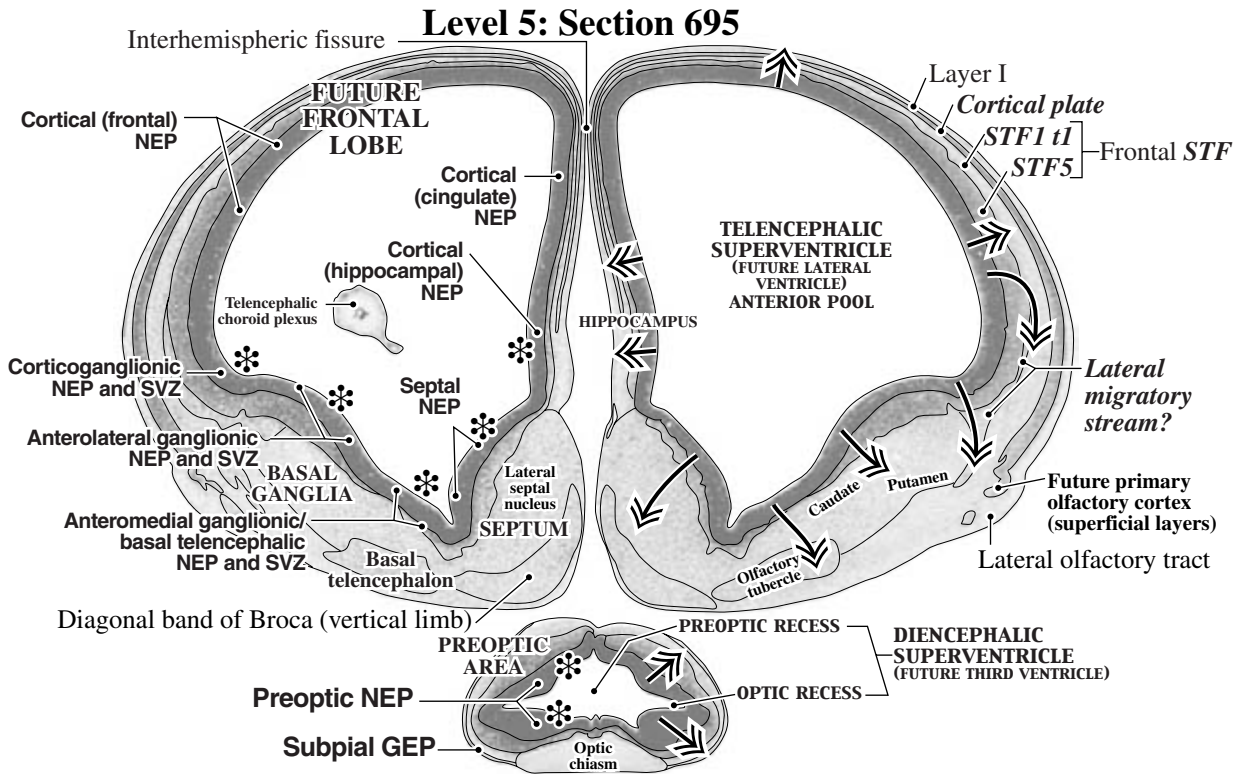
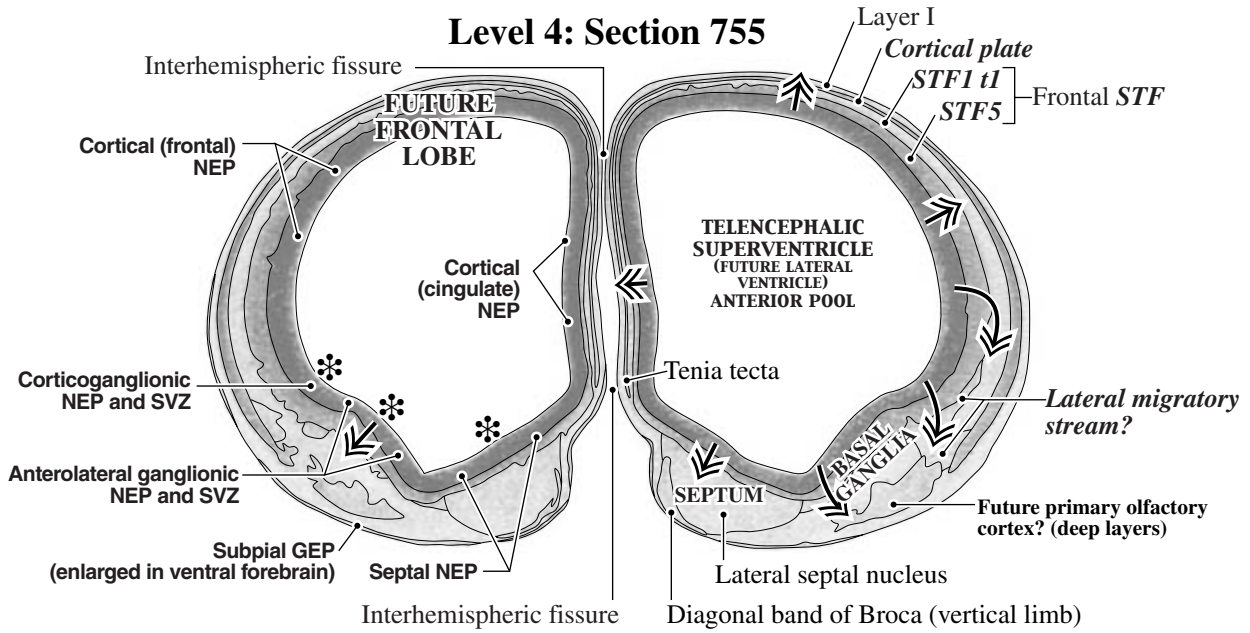
STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

Level 5: Section 695



2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

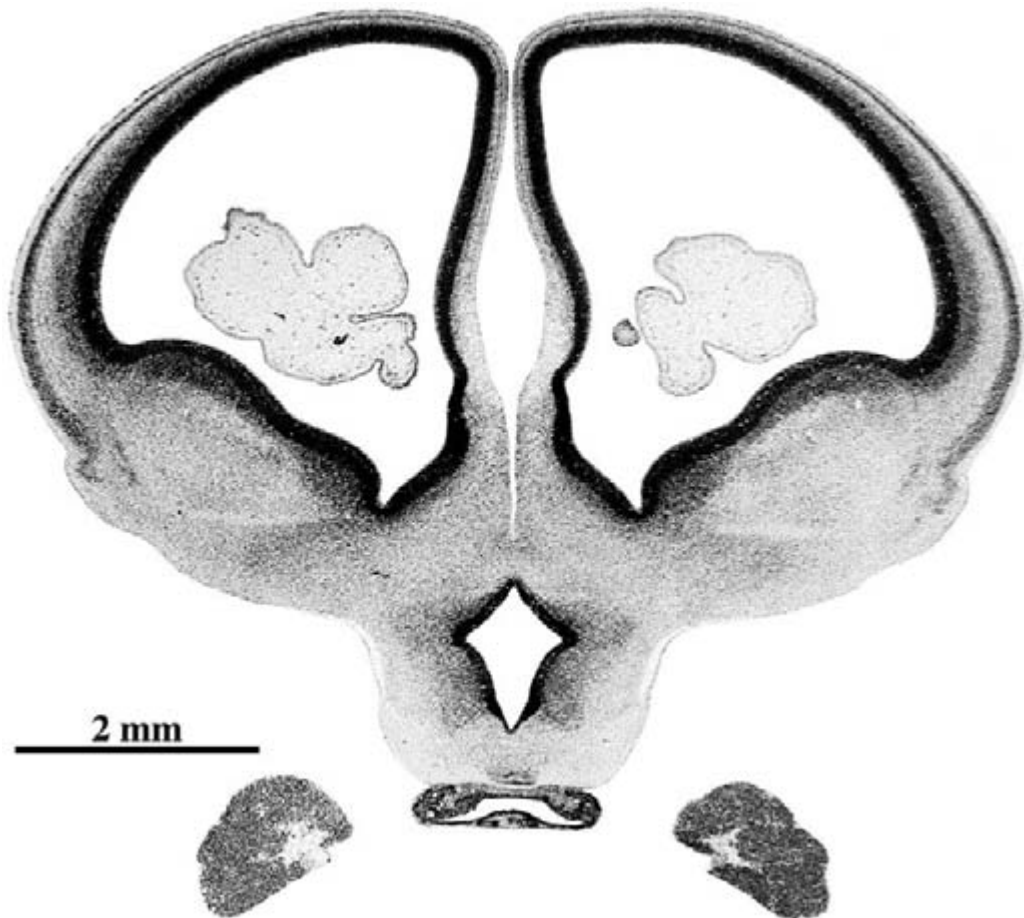
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 150A

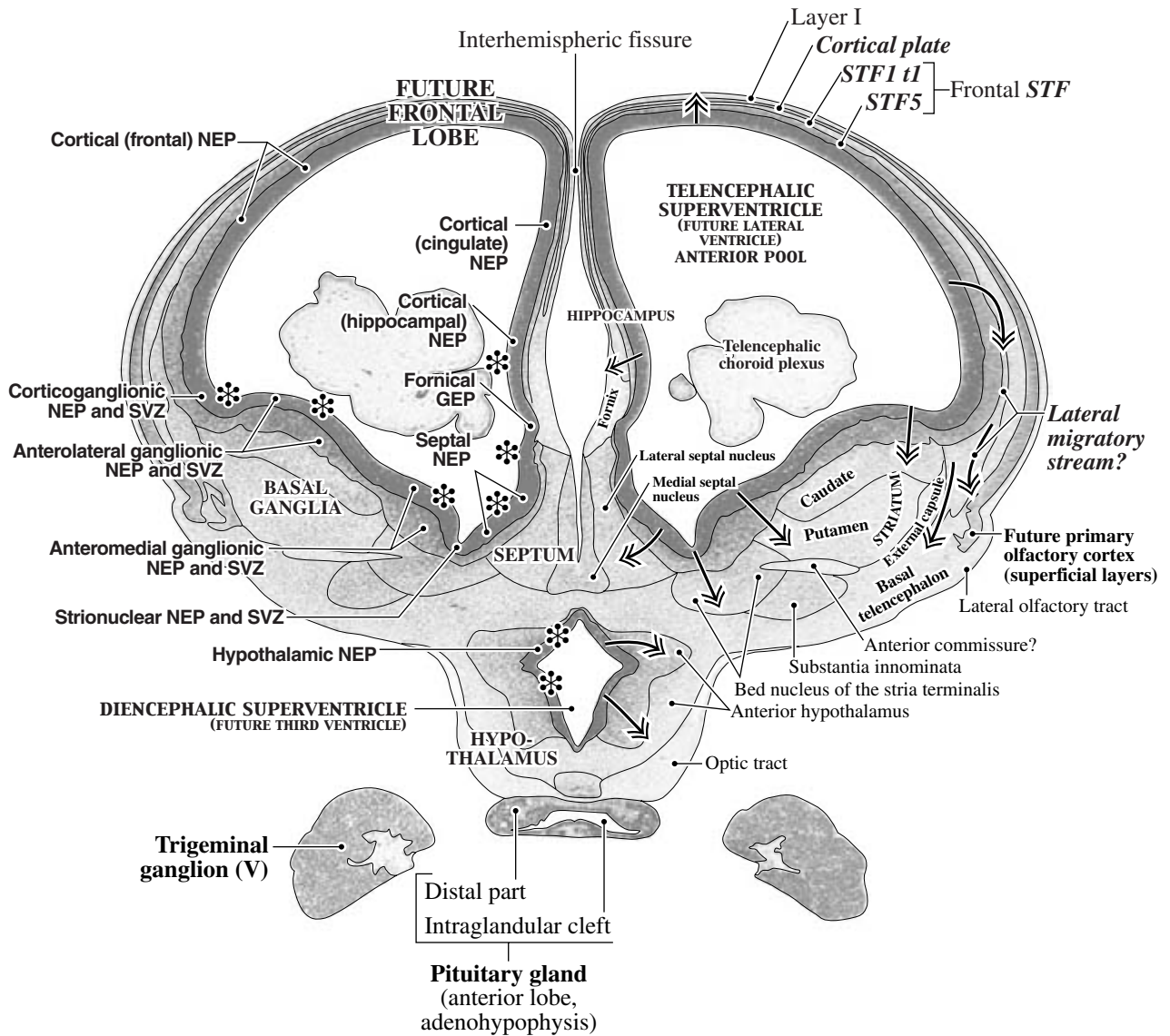
GW8 Coronal
CR 31 mm
C9226
Level 6: Section 653

**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*1I*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 151A

GW8 Coronal
CR 31 mm
C9226
Level 7: Section 635

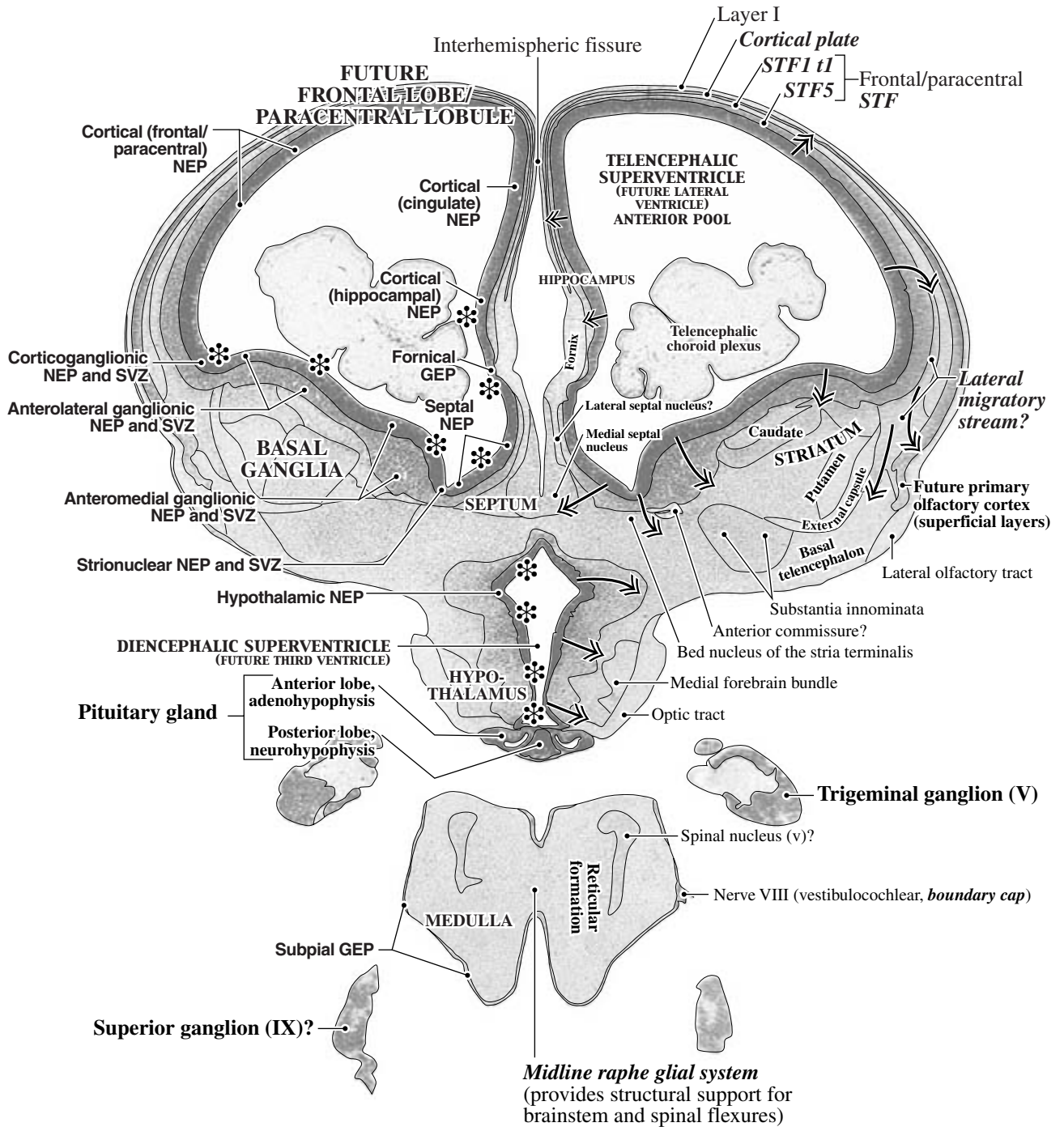
**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

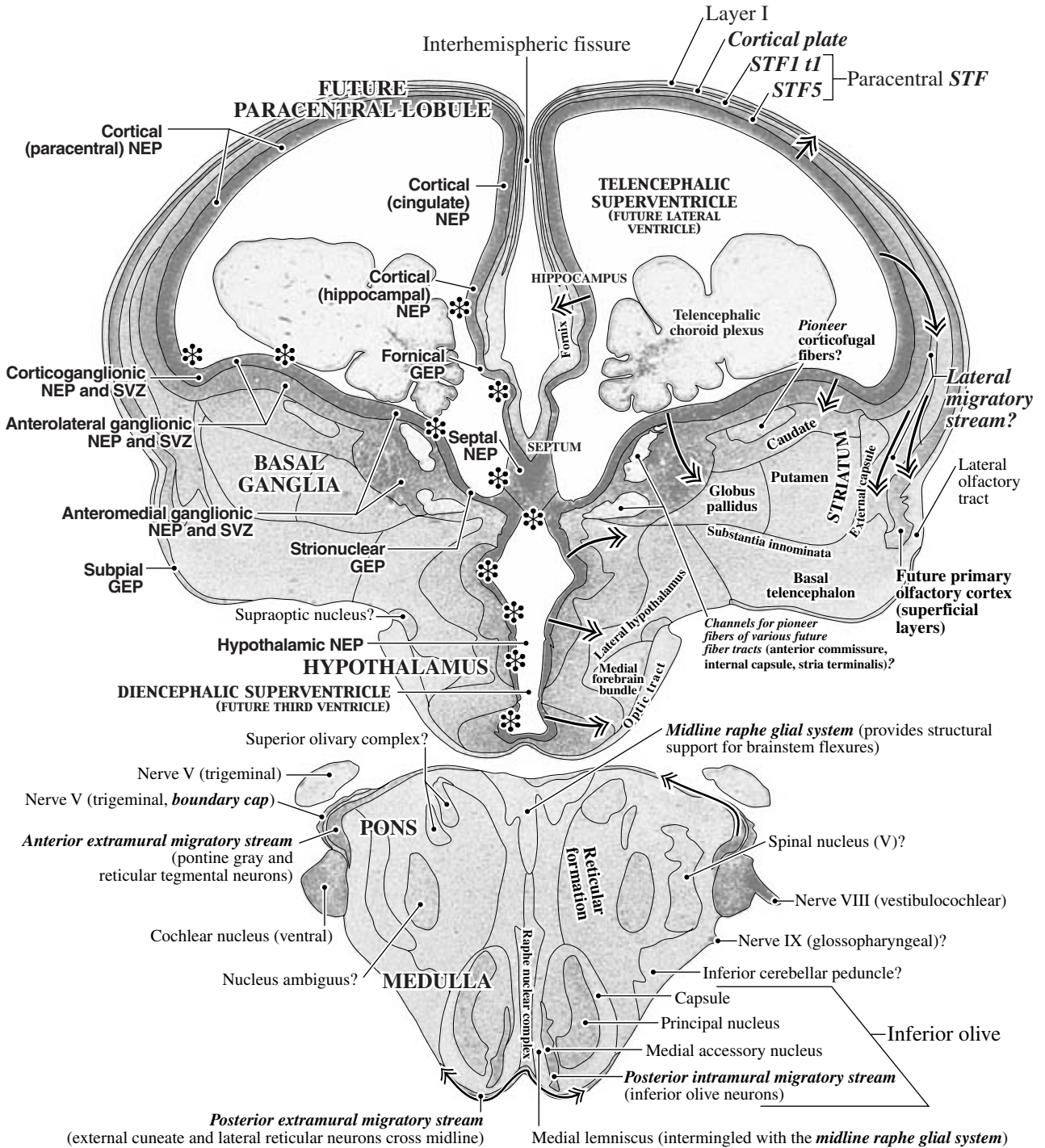
PLATE 152A**GW8 Coronal****CR 31 mm****C9226****Level 8: Section 605****LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

**2 mm**

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

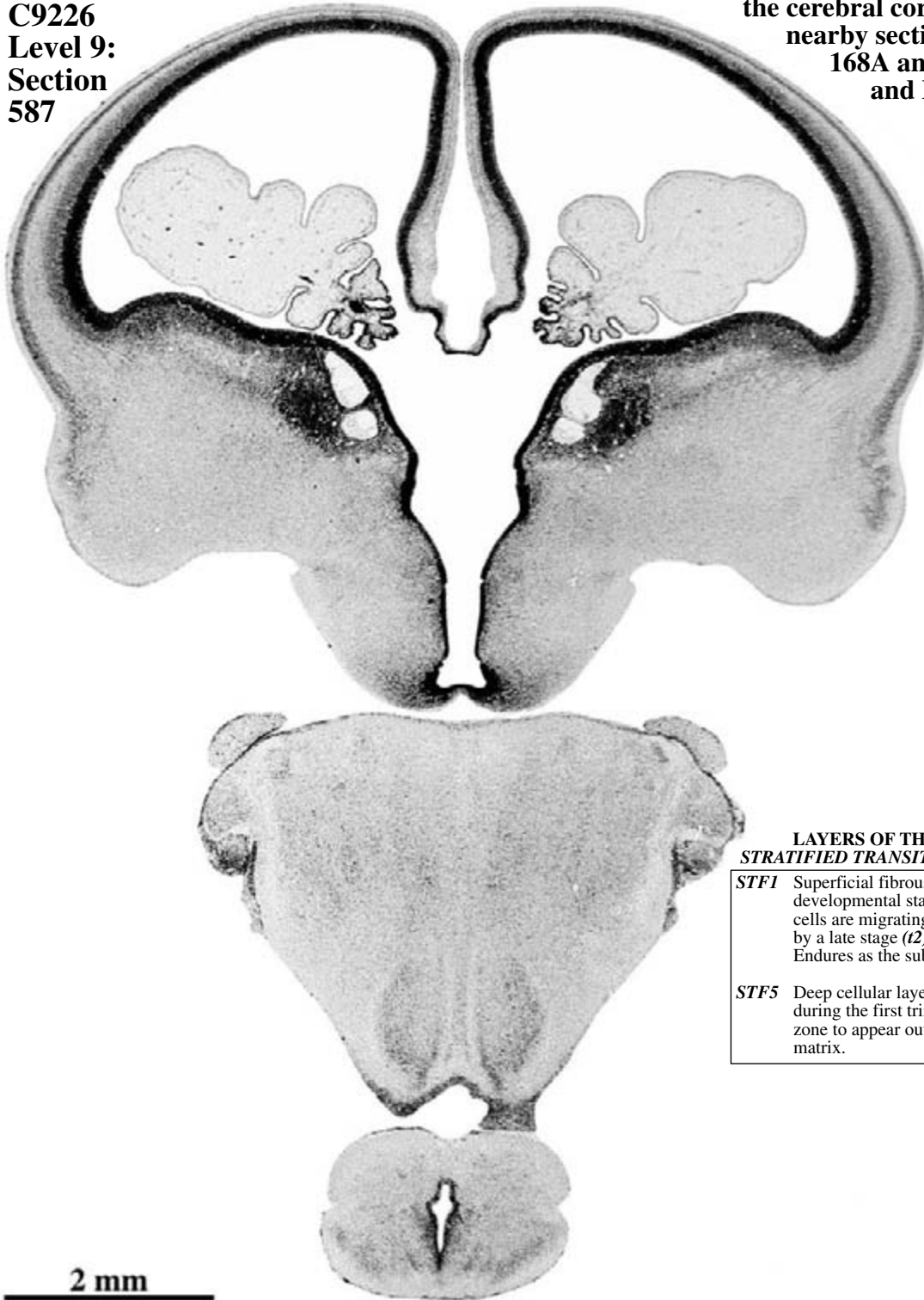
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 153A

GW8 Coronal
CR 31 mm
C9226
Level 9:
Section
587

See high-magnification views of
the cerebral cortex from
nearby sections in Plates
168A and B to 169A
and B.

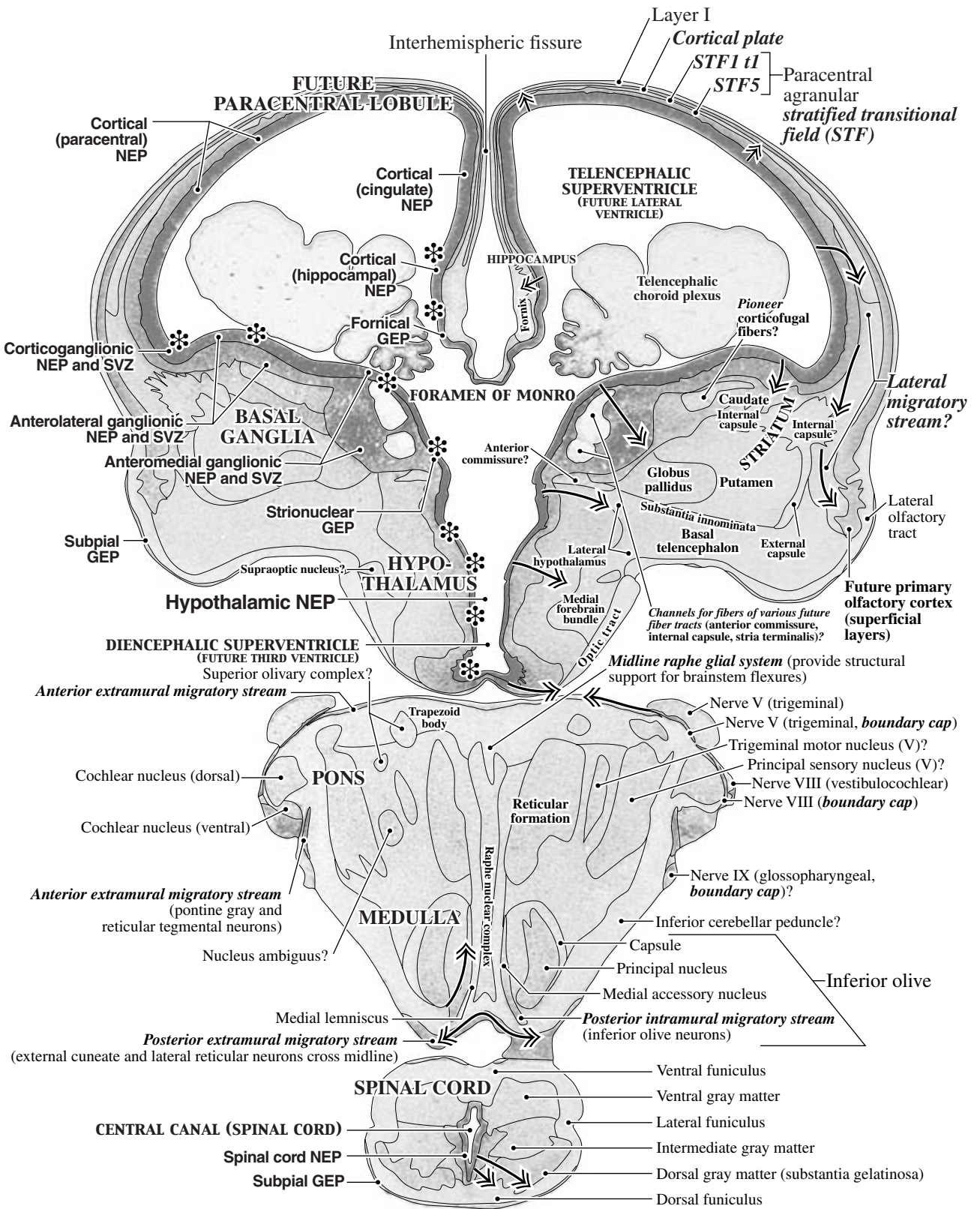


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1*** Superficial fibrous layer with an early developmental stage (***t1***) when many cells are migrating through it, followed by a late stage (***t2***) with sparse cells. Endures as the subcortical white matter.
- STF5*** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

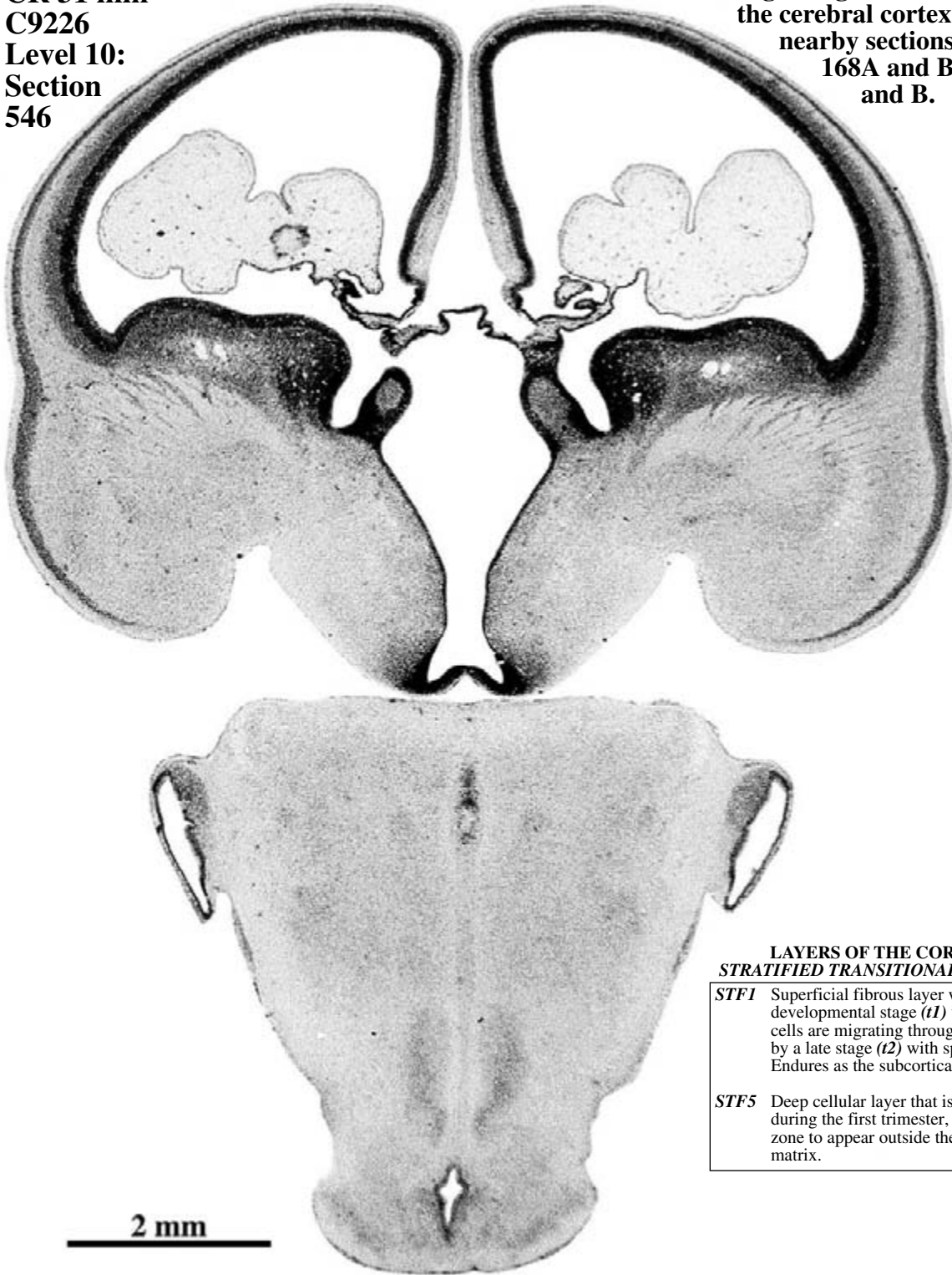
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 154A

GW8 Coronal
CR 31 mm
C9226
Level 10:
Section
546

See high-magnification views of
the cerebral cortex from
nearby sections in Plates
168A and B to 169A
and B.

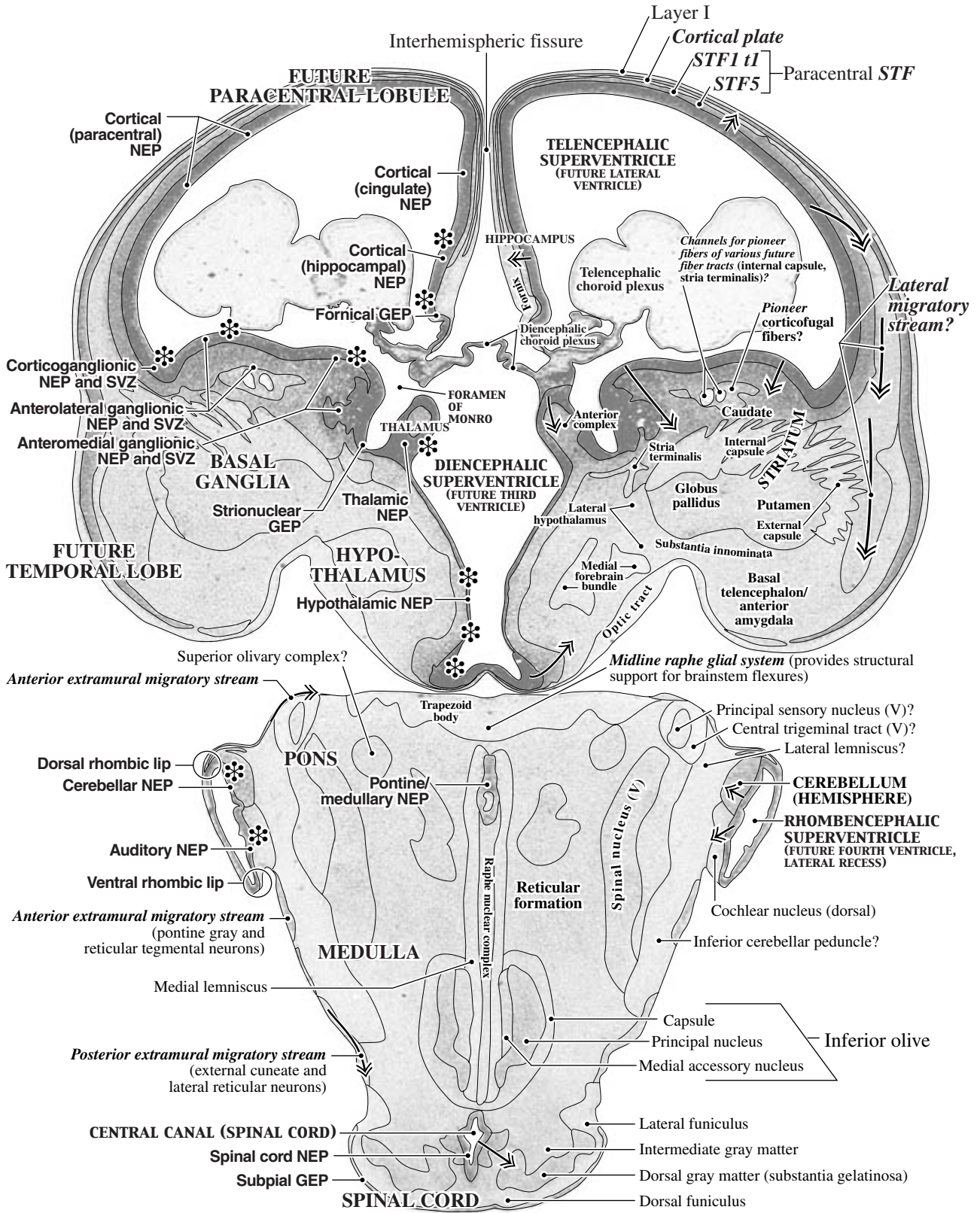


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



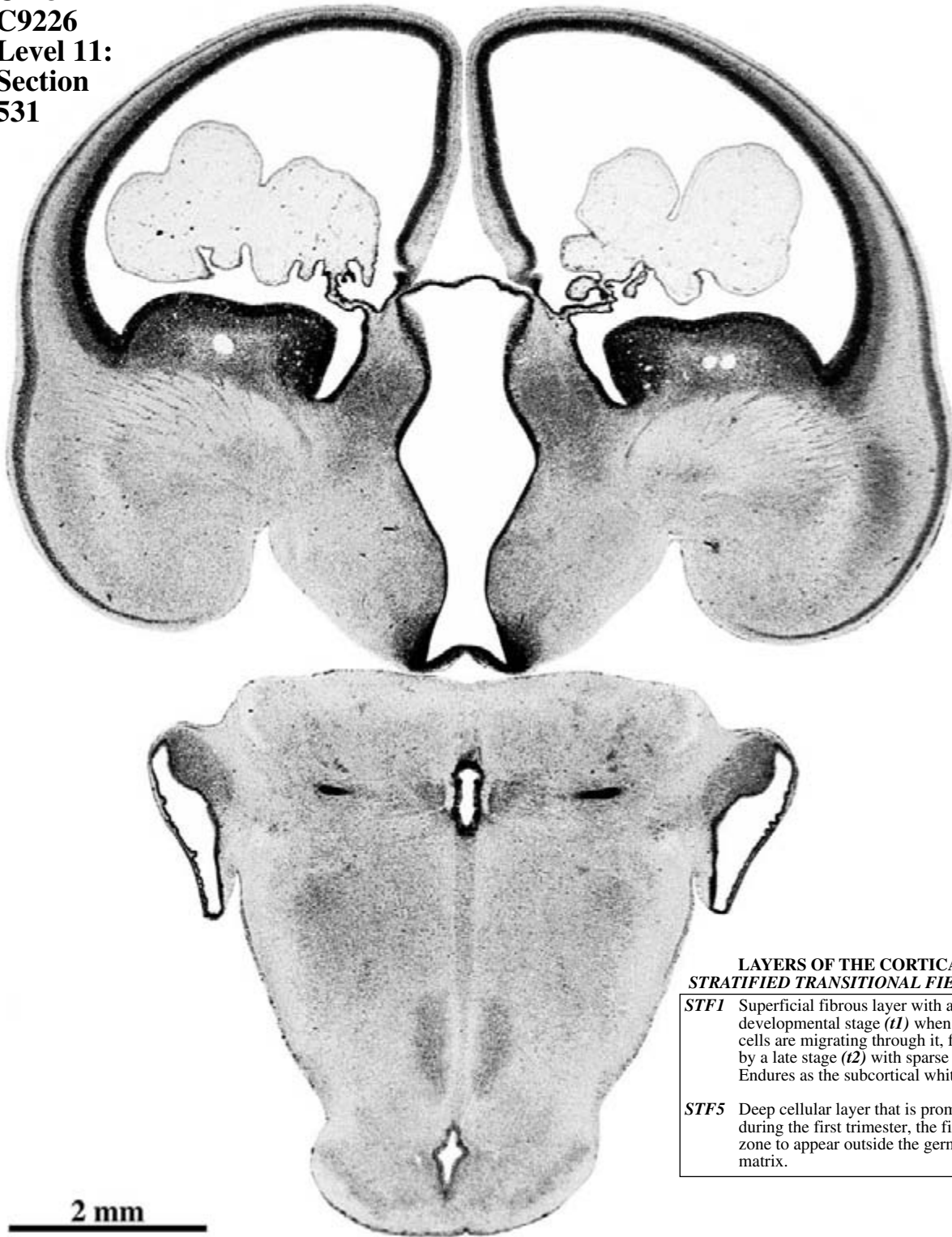
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 155A

GW8 Coronal
CR 31 mm
C9226
Level 11:
Section
531

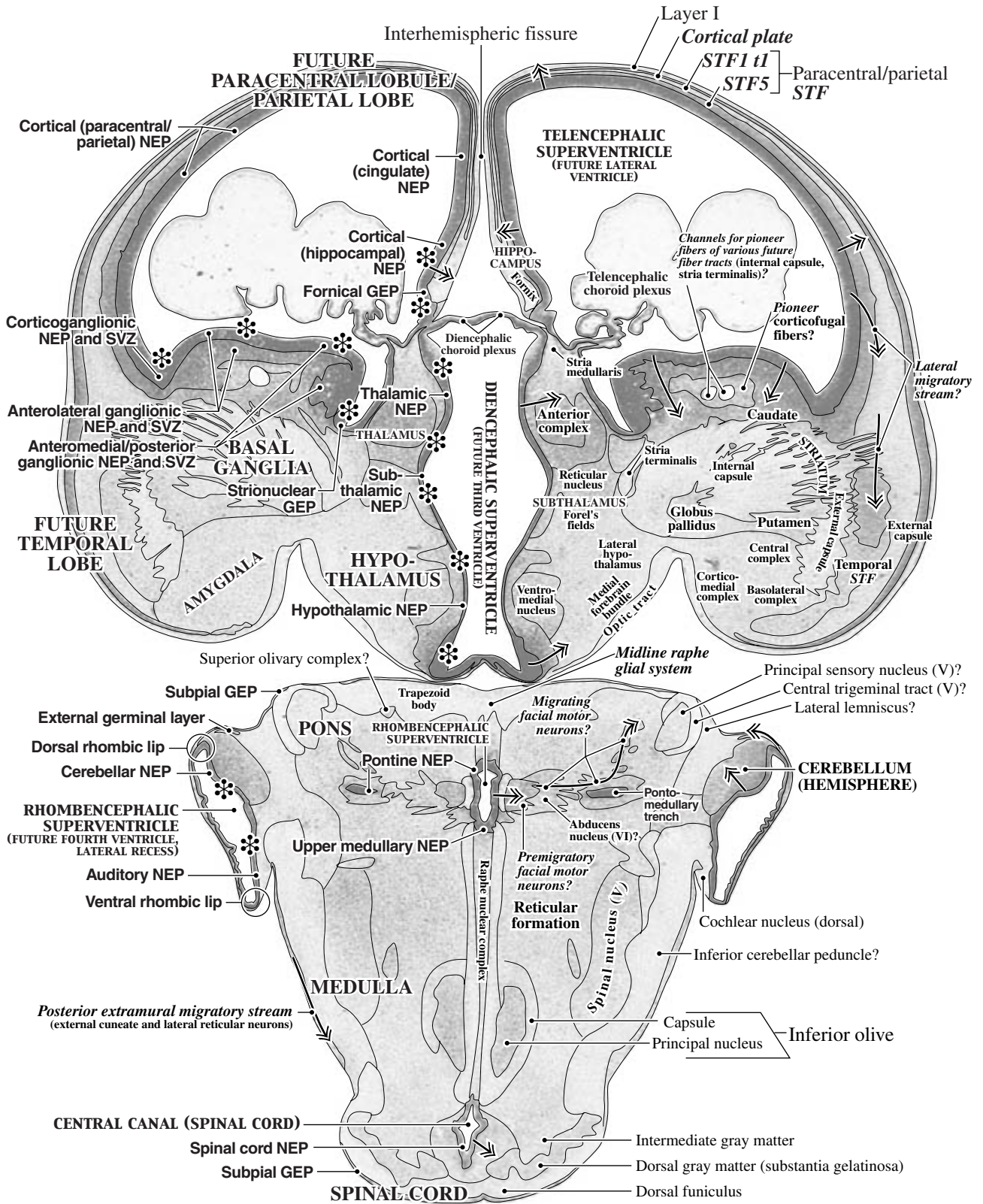


**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

- STF1*** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5*** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

2 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



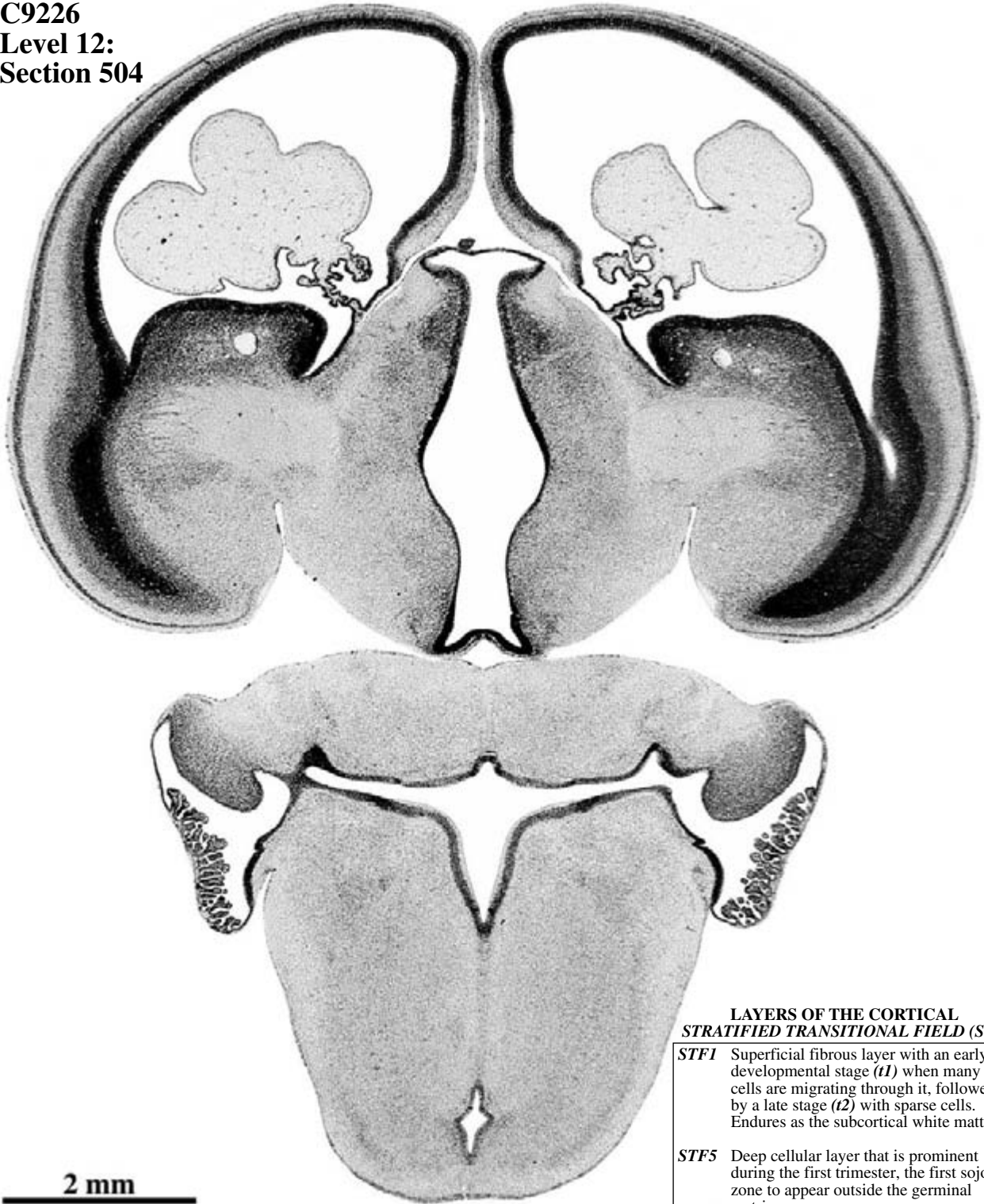
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 156A

GW8 Coronal
CR 31 mm
C9226
Level 12:
Section 504



**LAYERS OF THE CORTICAL
STRATIFIED TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

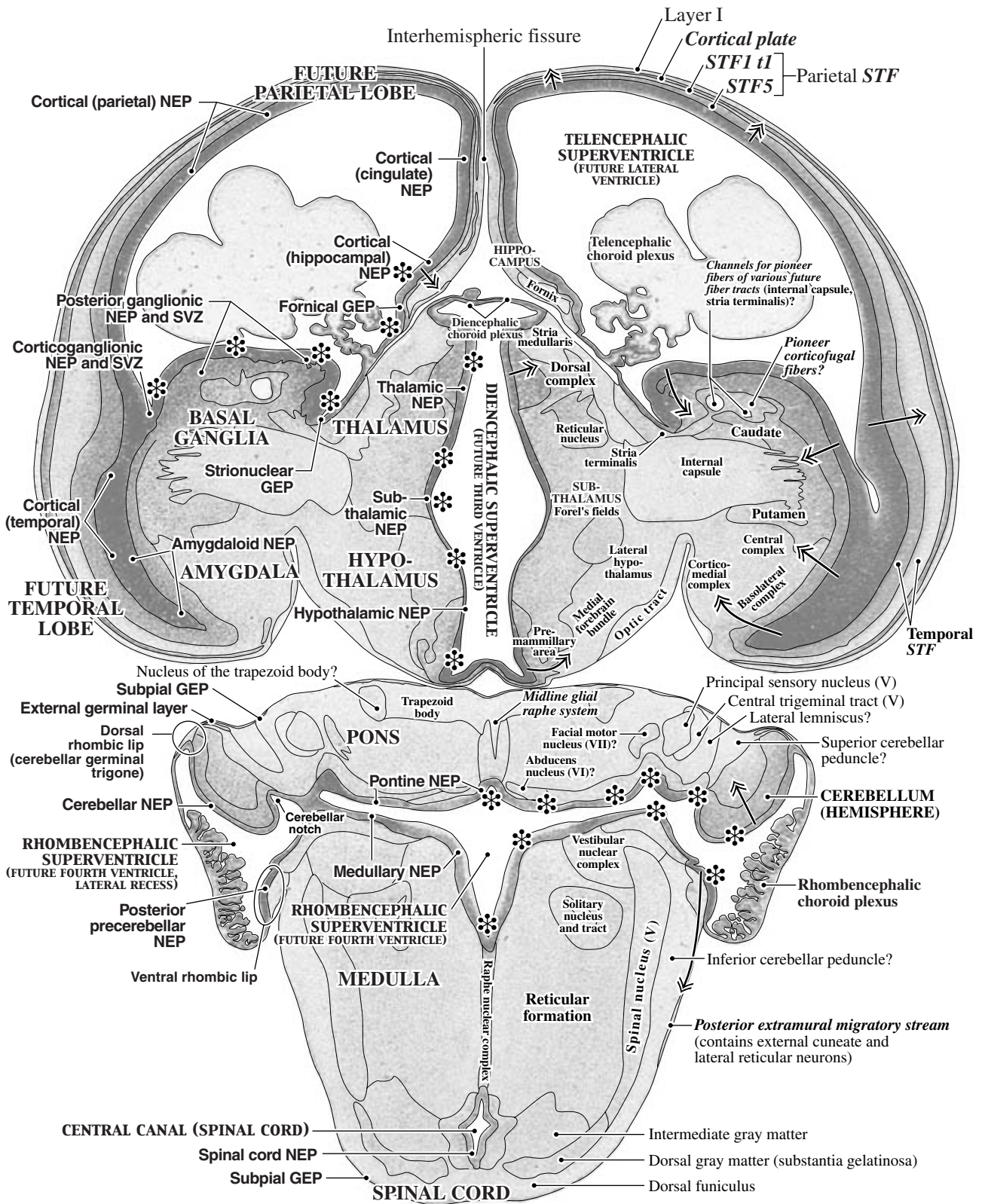
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**



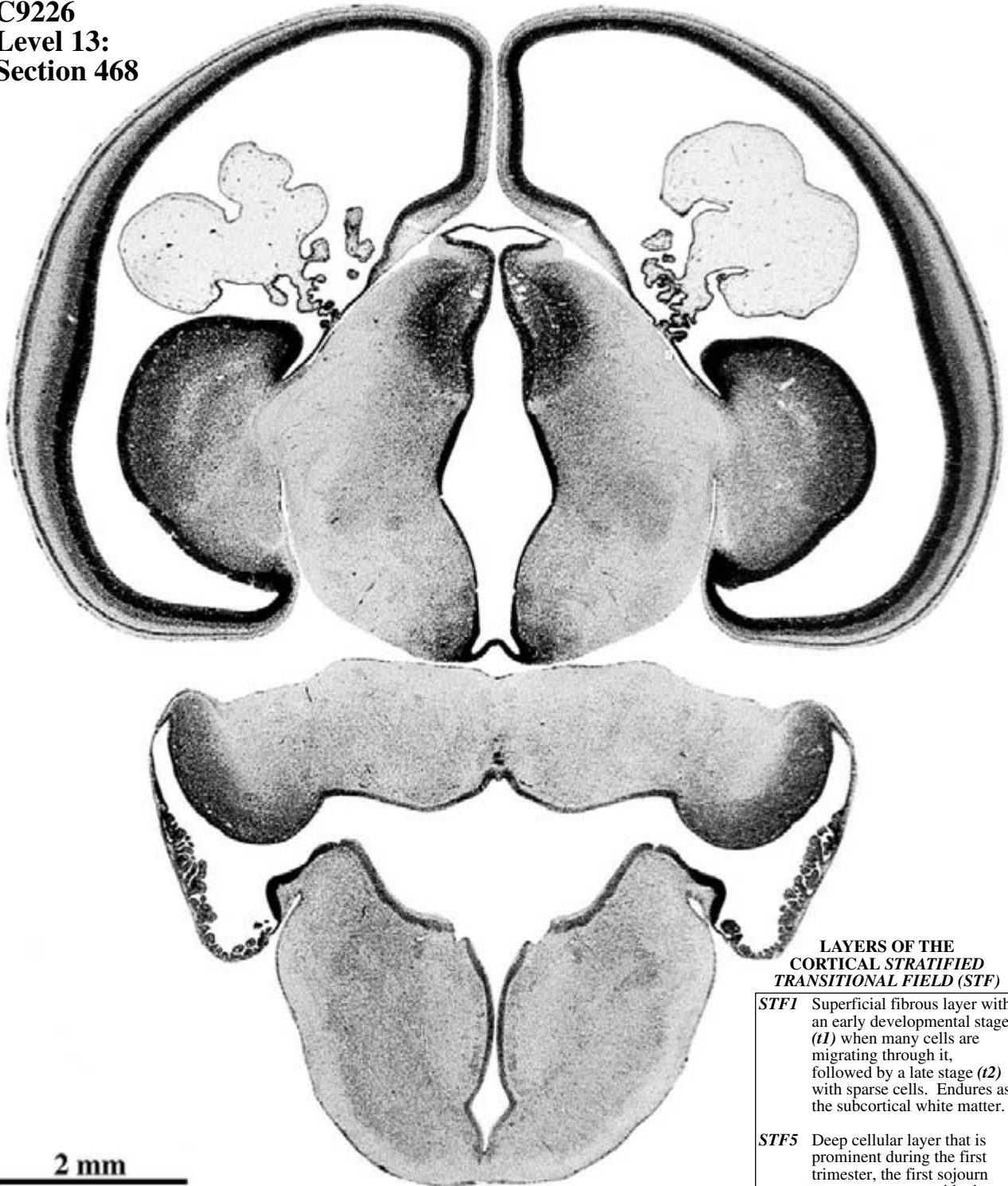
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 157A

GW8 Coronal
CR 31 mm
C9226
Level 13:
Section 468



**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

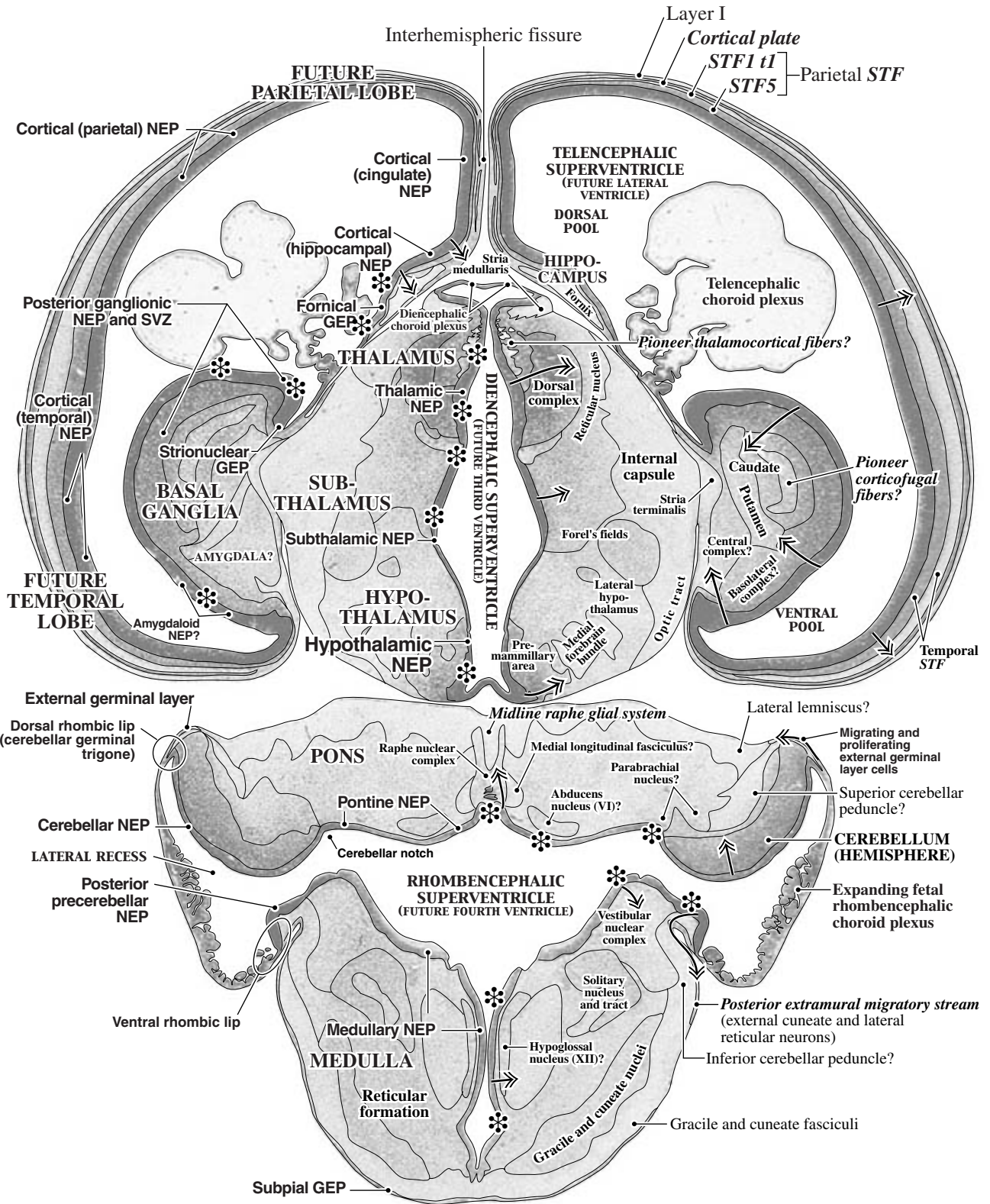
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or Bold



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 158A

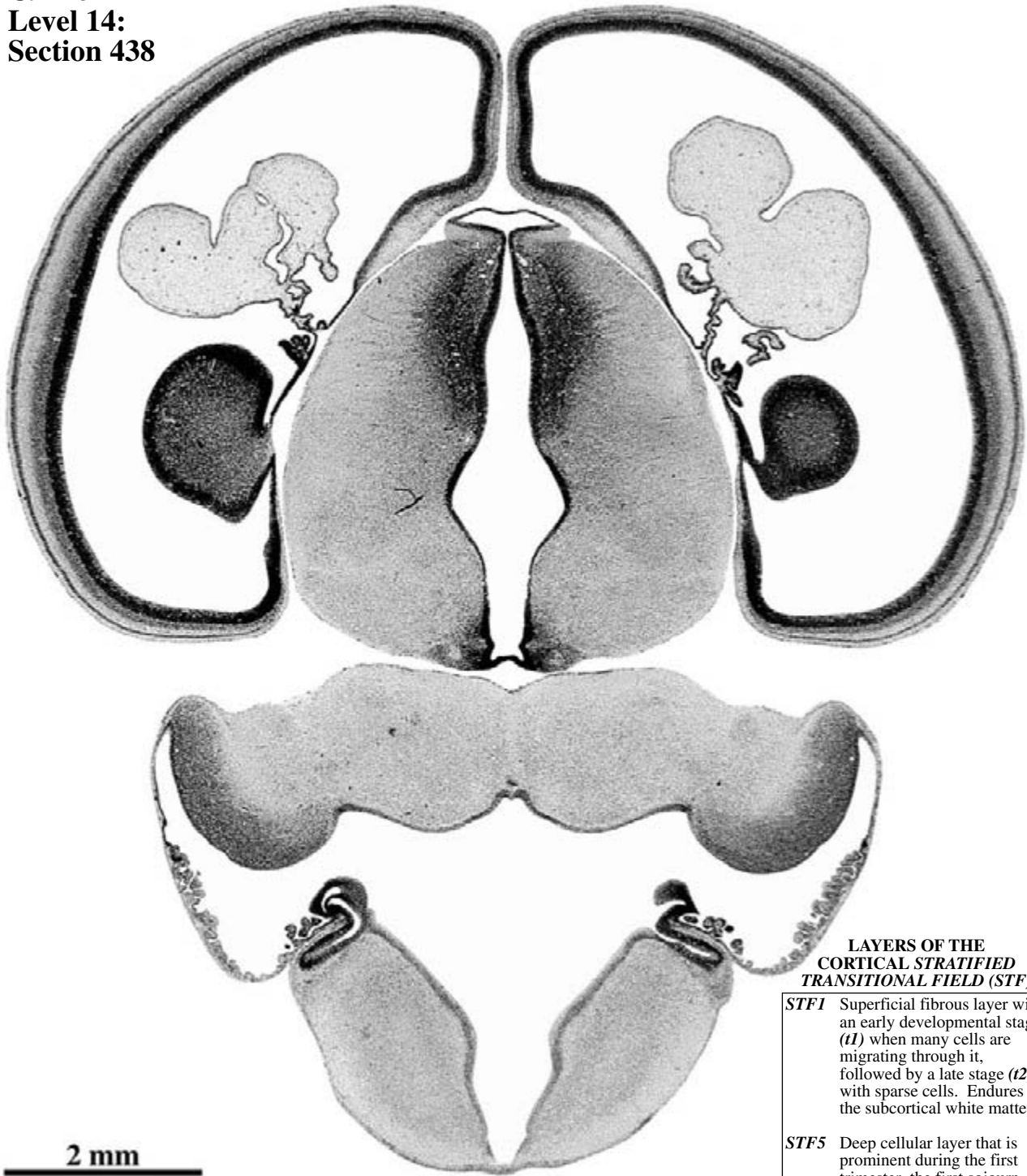
GW8 Coronal

CR 31 mm

C9226

Level 14:

Section 438

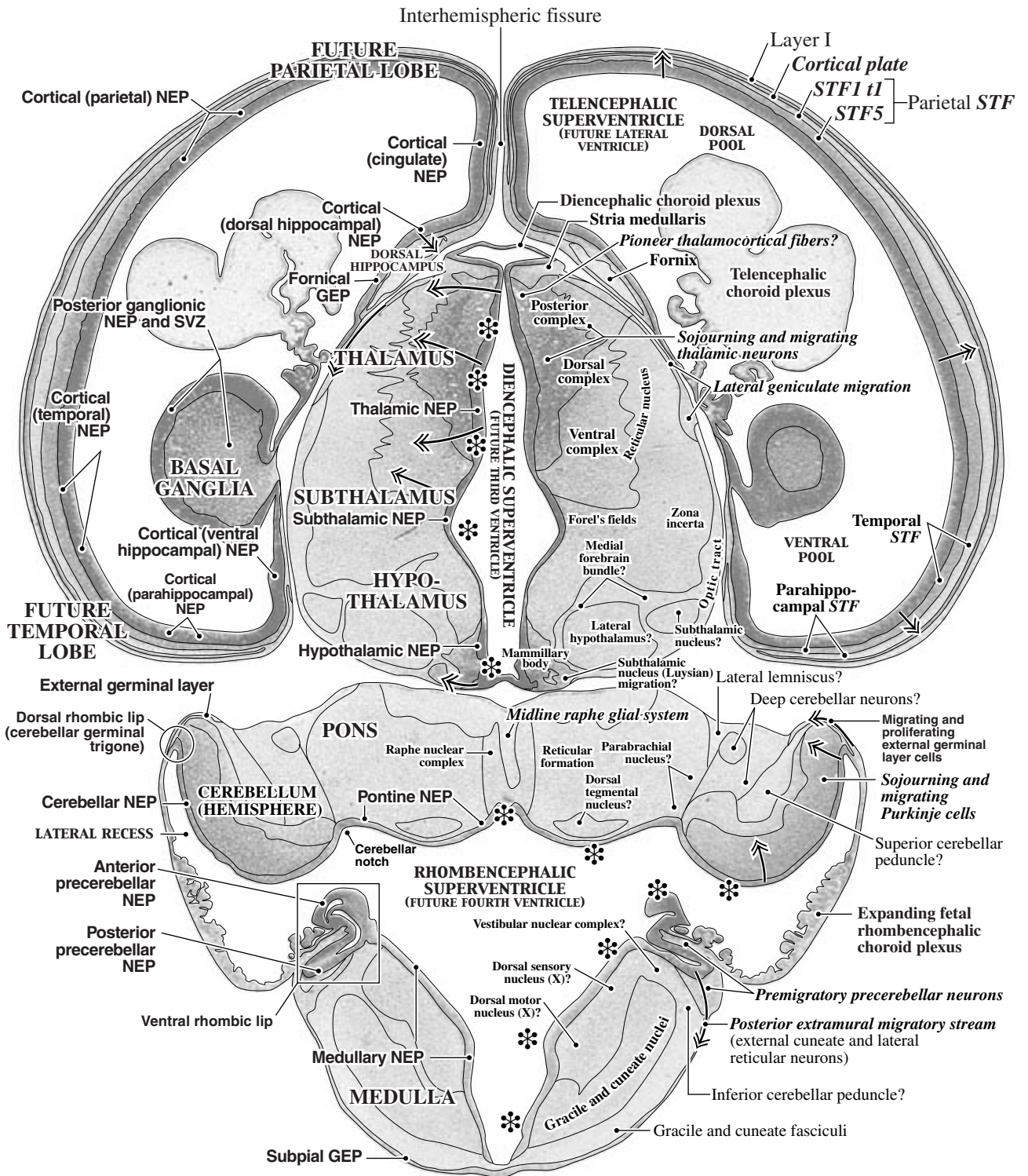


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



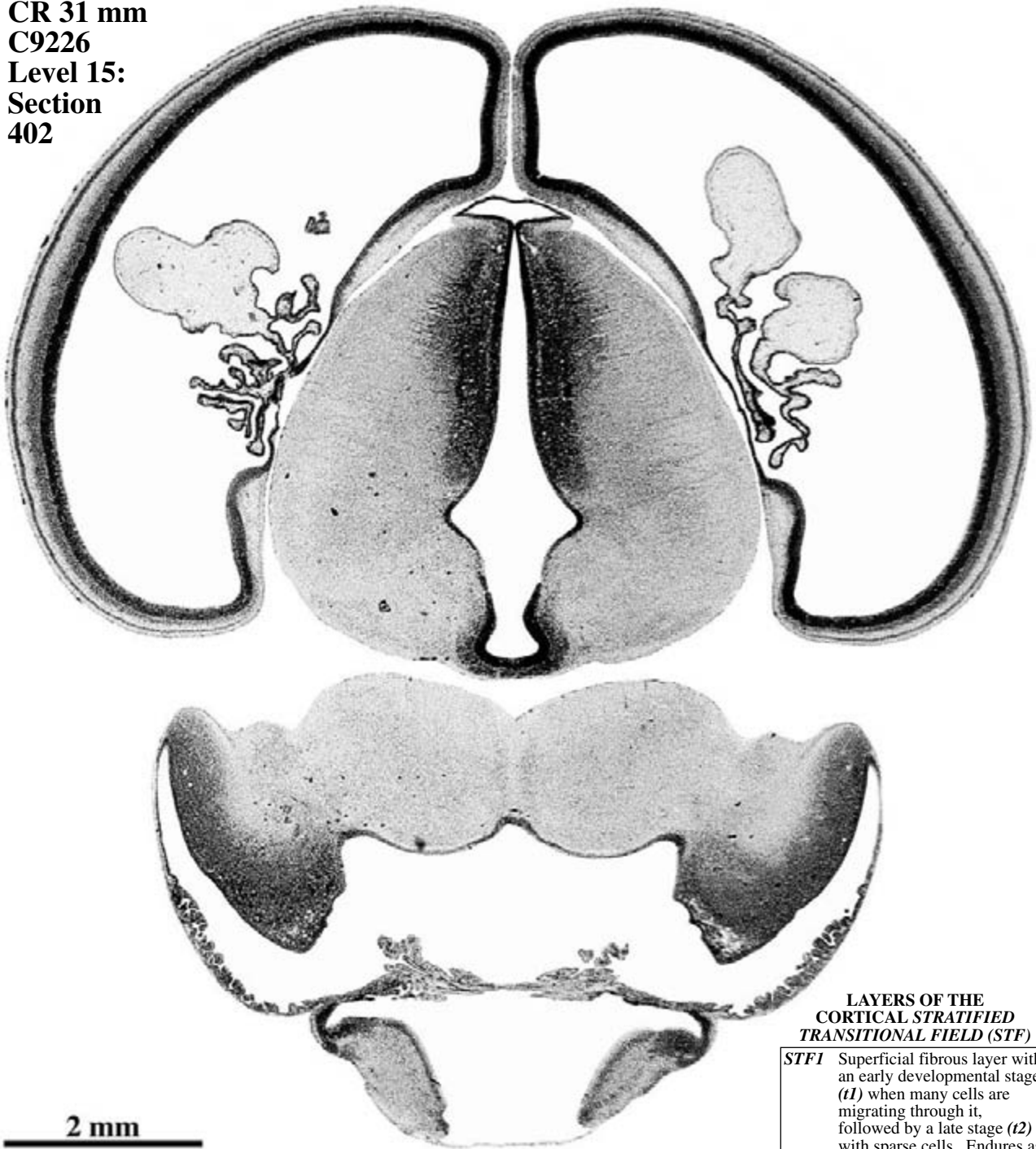
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 159A

GW8 Coronal
CR 31 mm
C9226
Level 15:
Section
402



**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

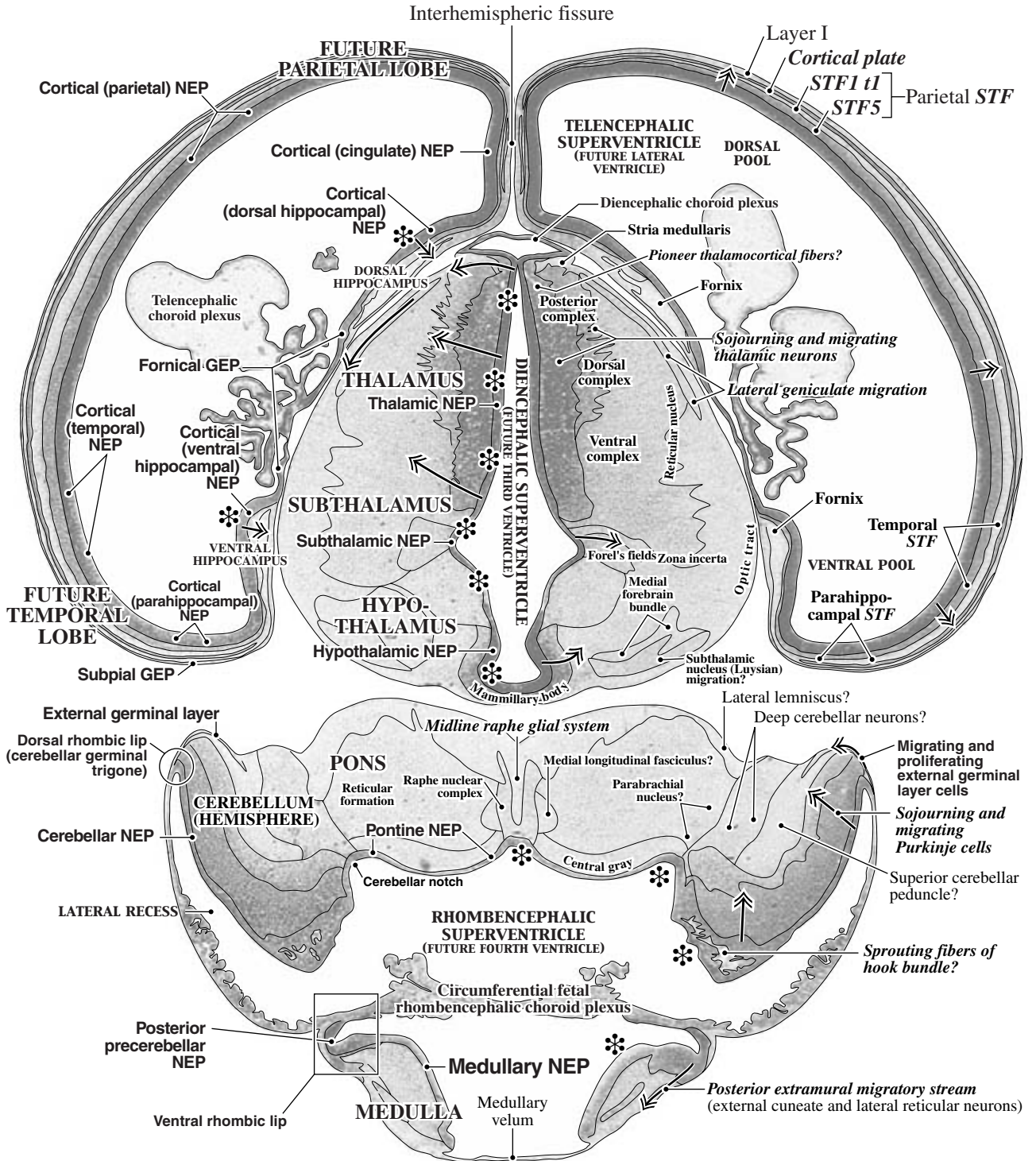
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**



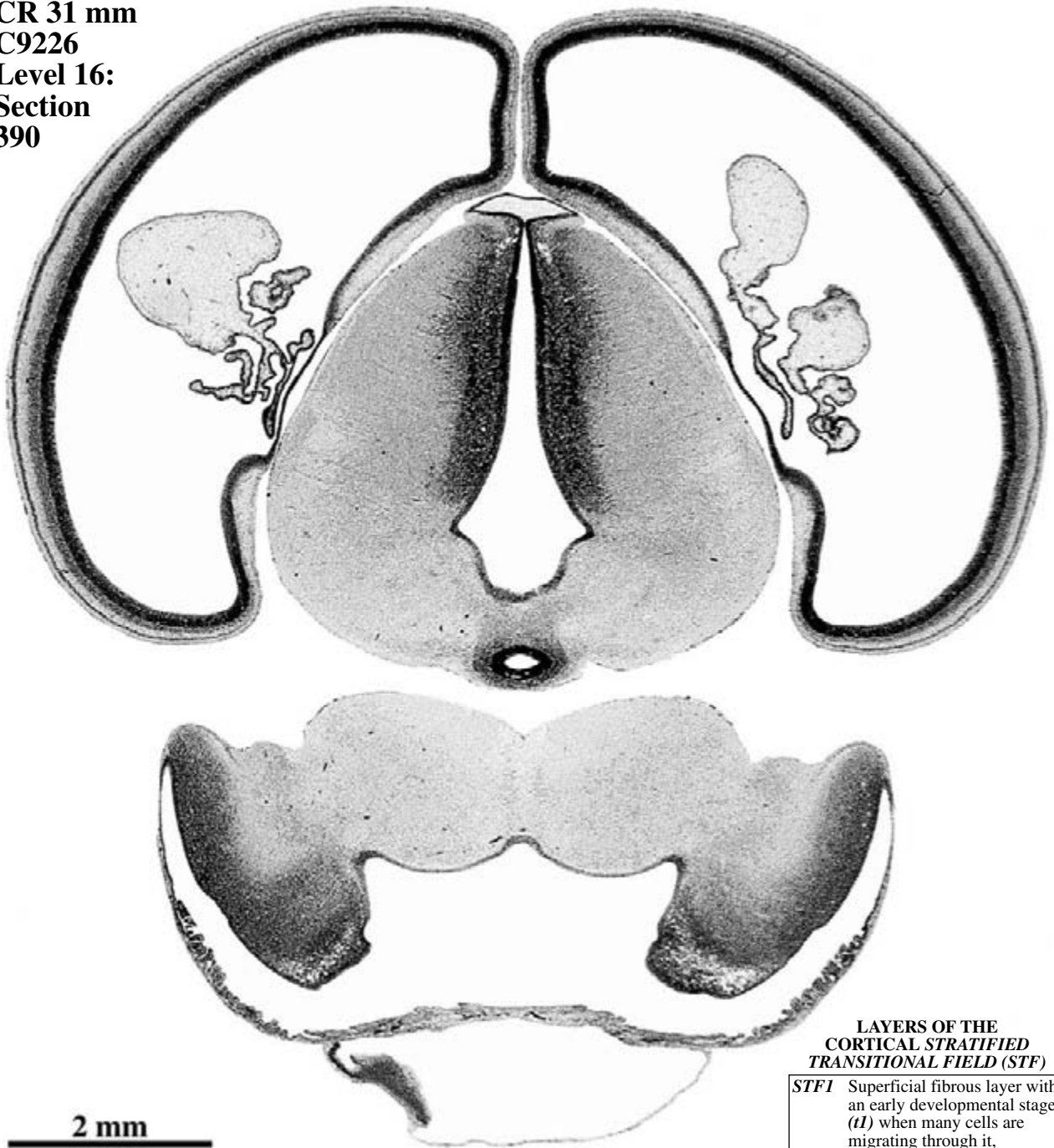
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 160A

GW8 Coronal
CR 31 mm
C9226
Level 16:
Section
390

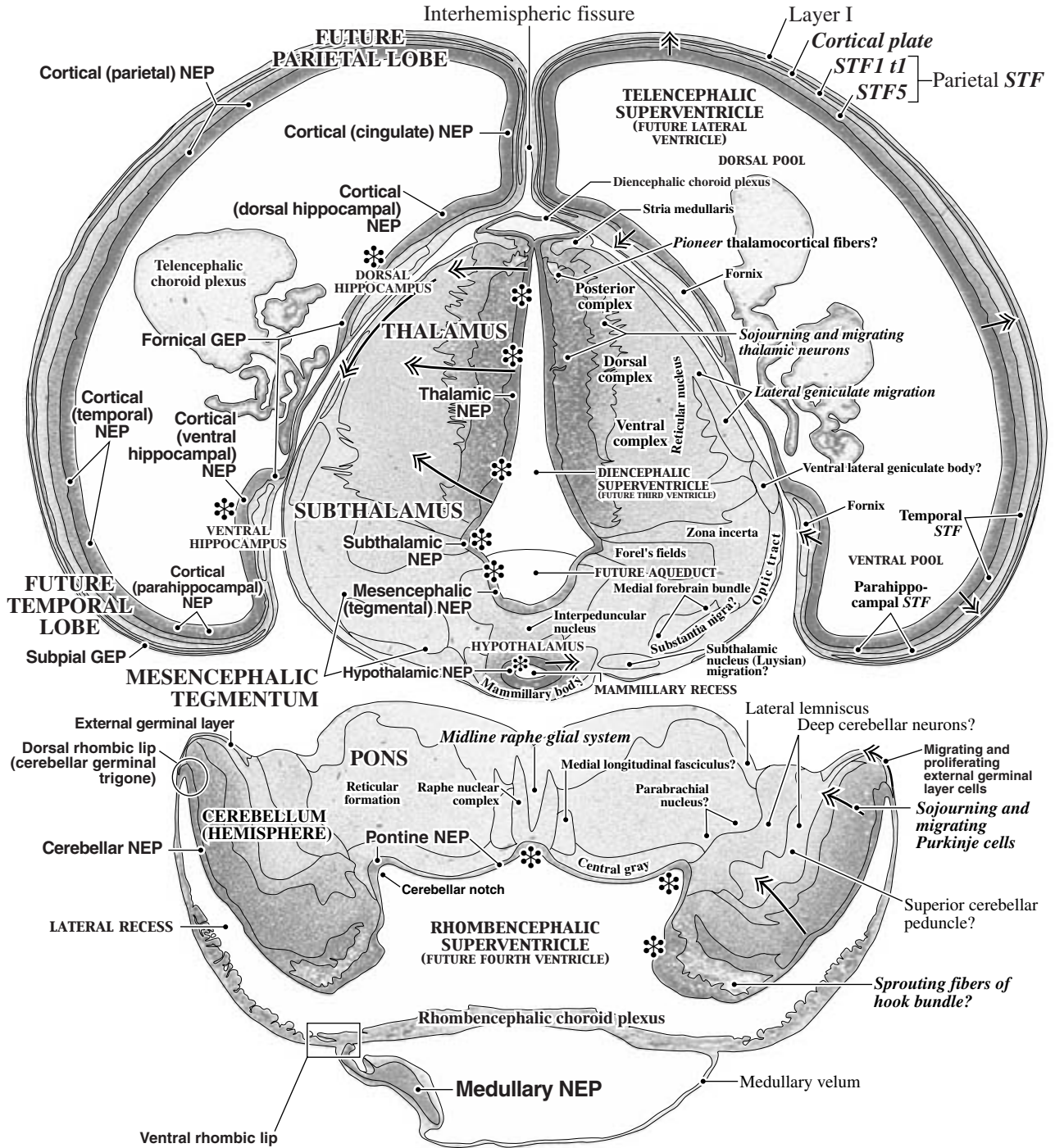


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (***11***) when many cells are migrating through it, followed by a late stage (***12***) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



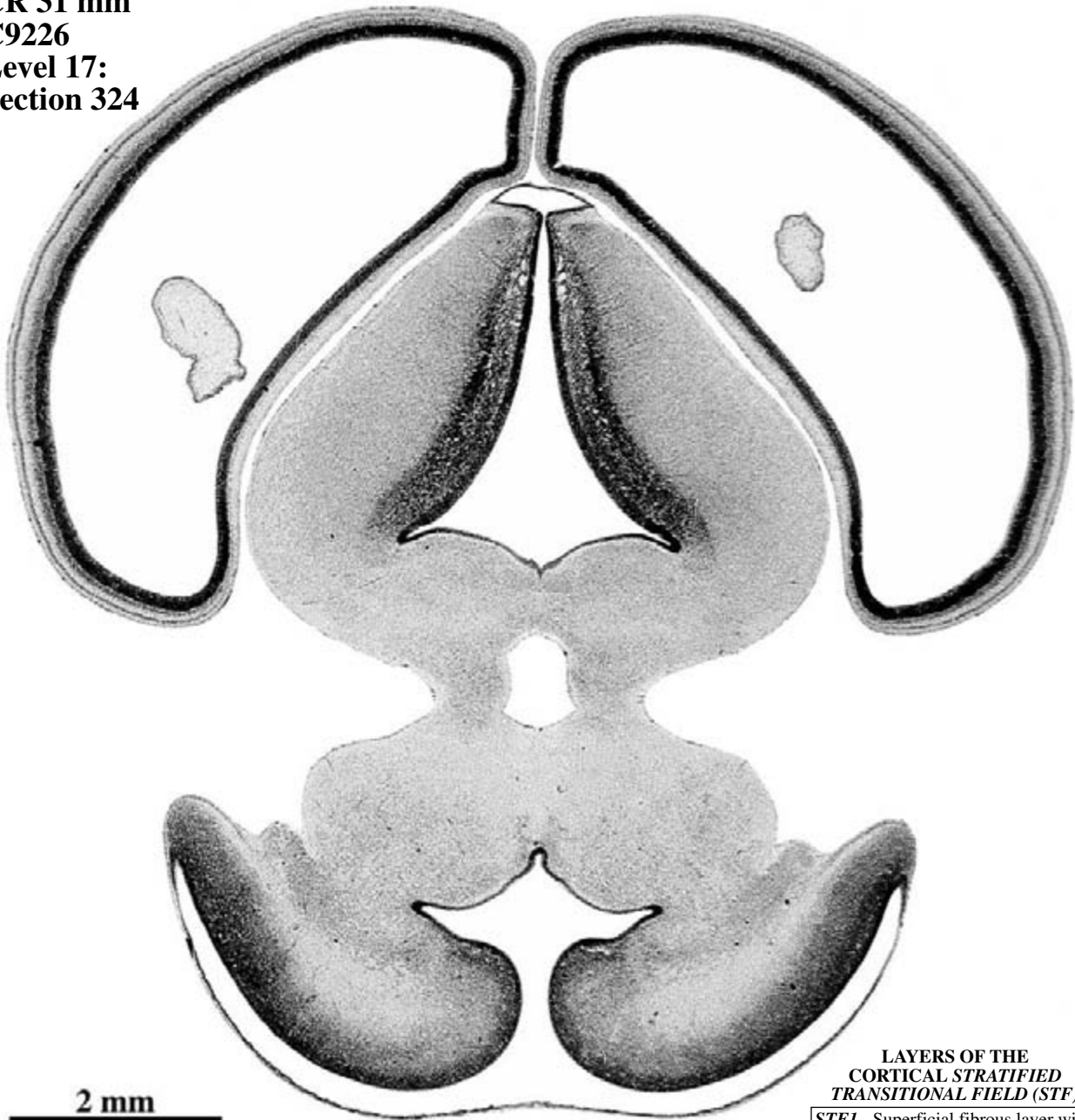
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 161A

GW8 Coronal
CR 31 mm
C9226
Level 17:
Section 324

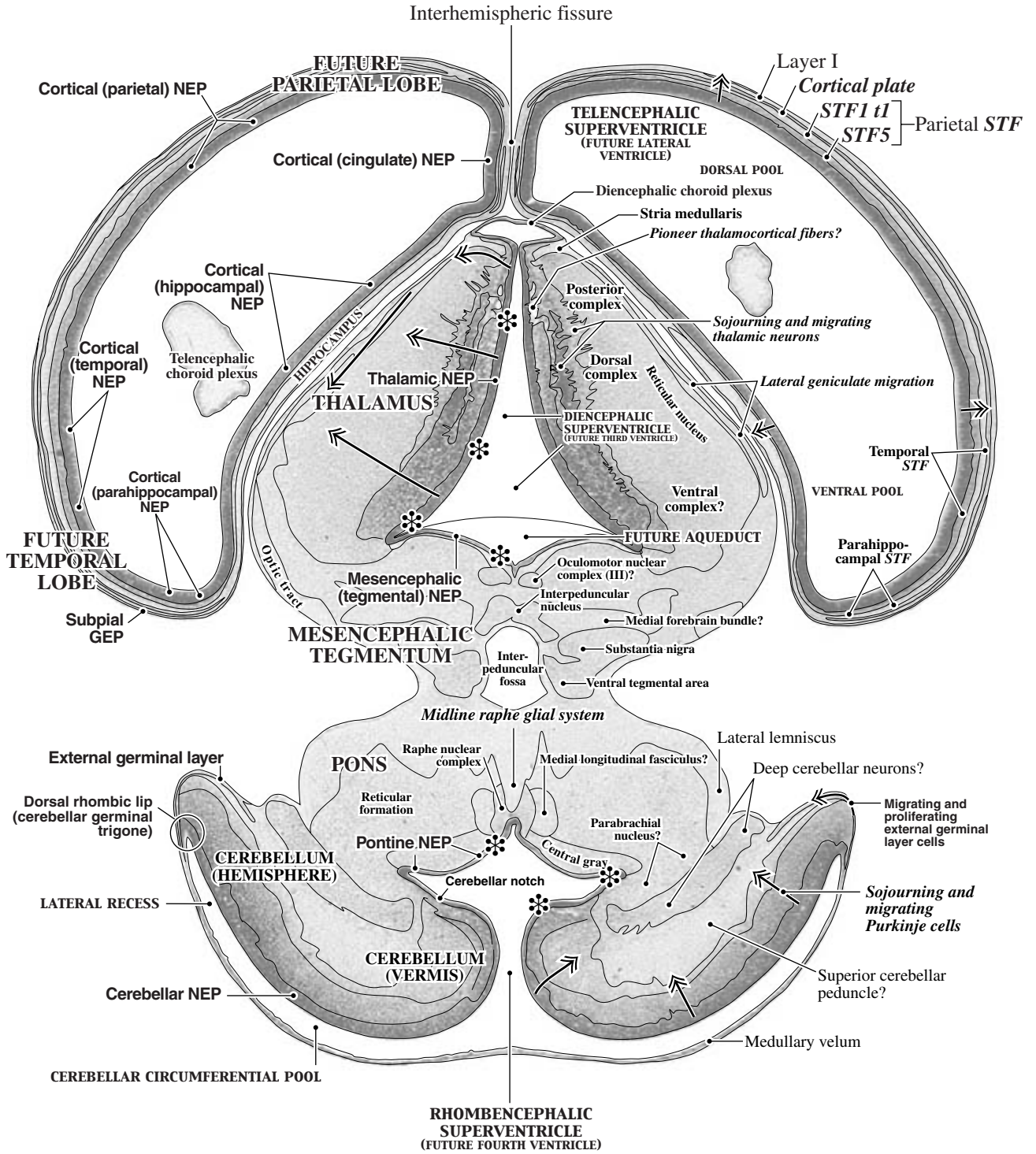


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



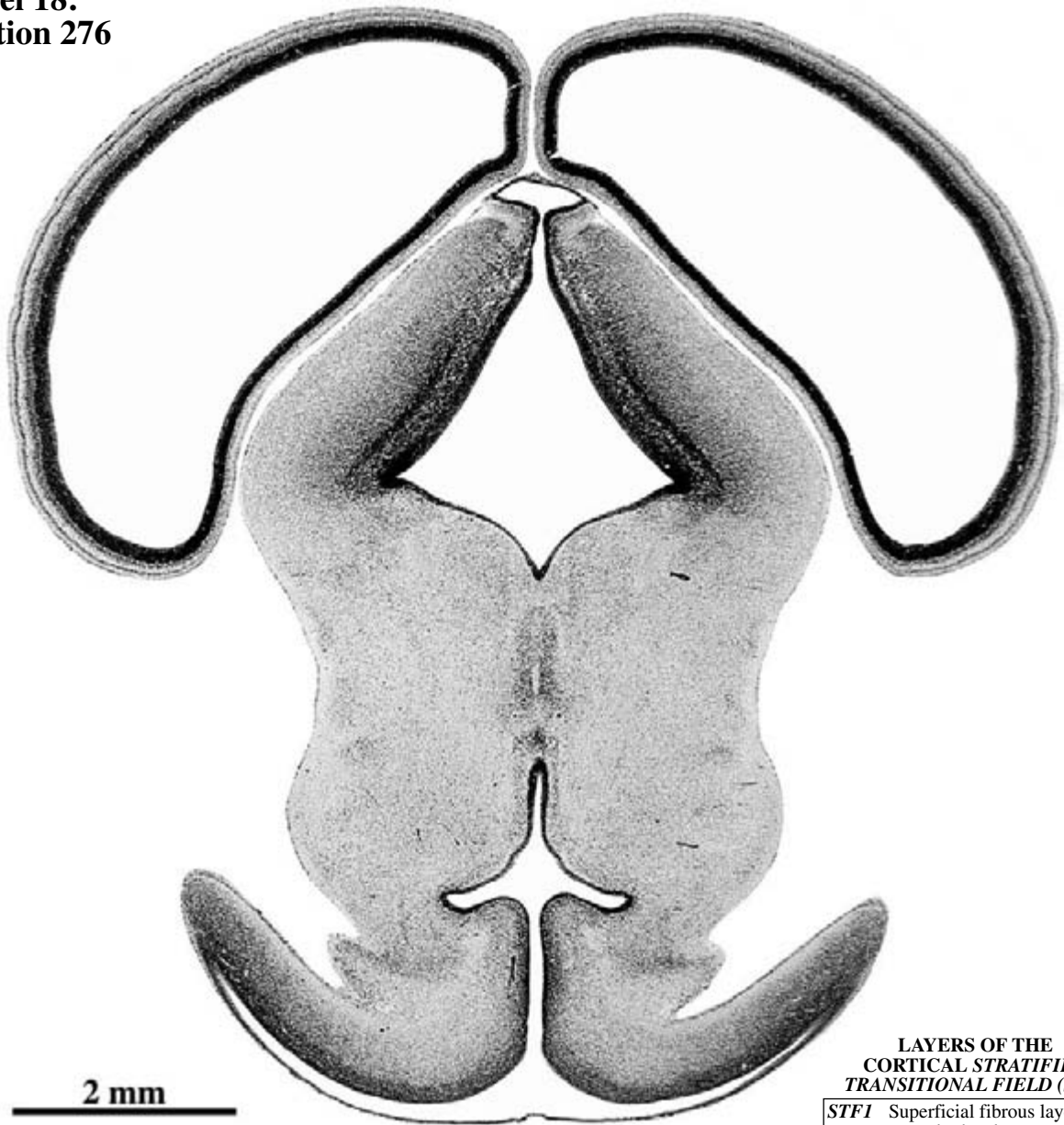
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 162A

GW8 Coronal
CR 31 mm
C9226
Level 18:
Section 276

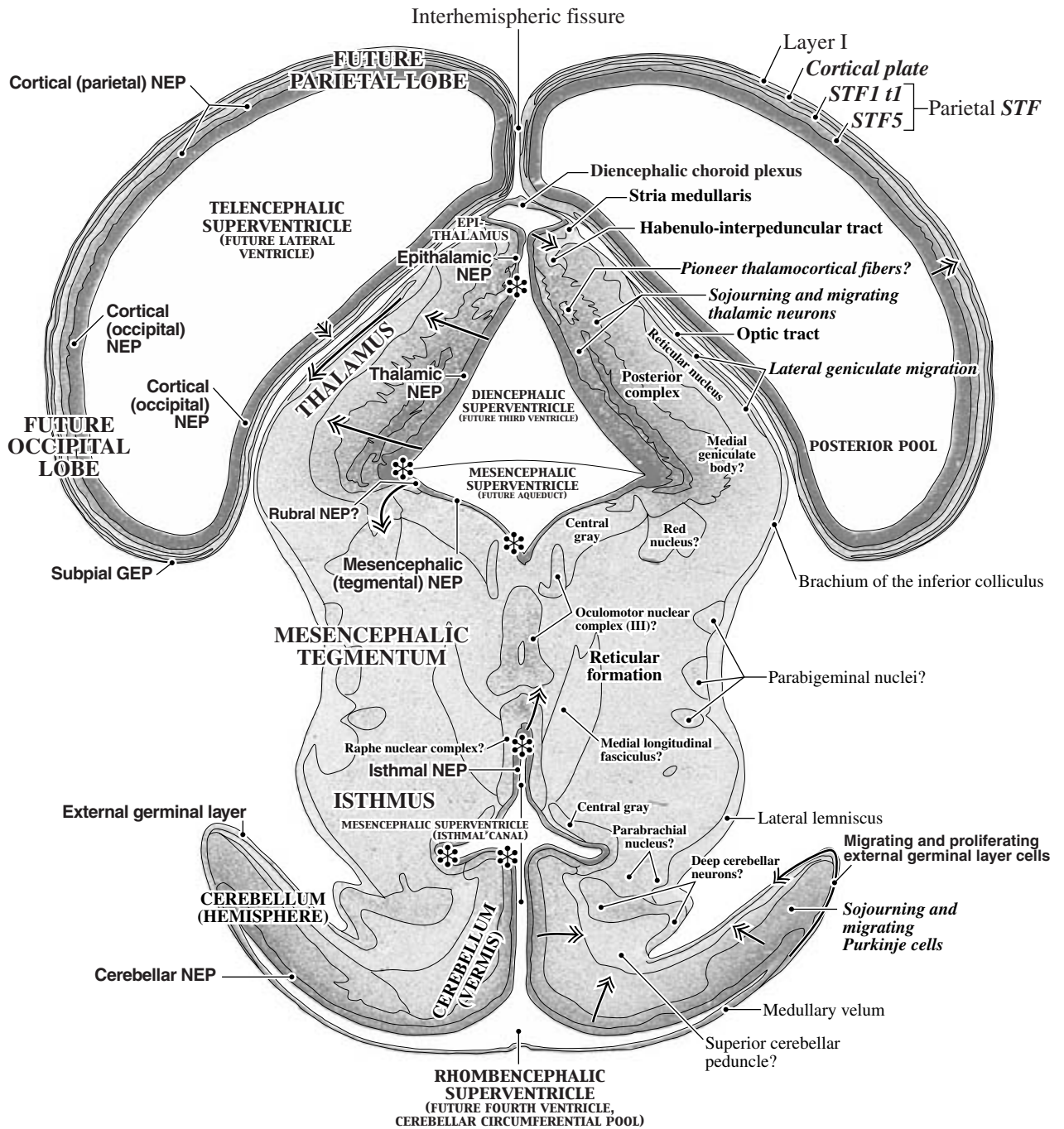


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold



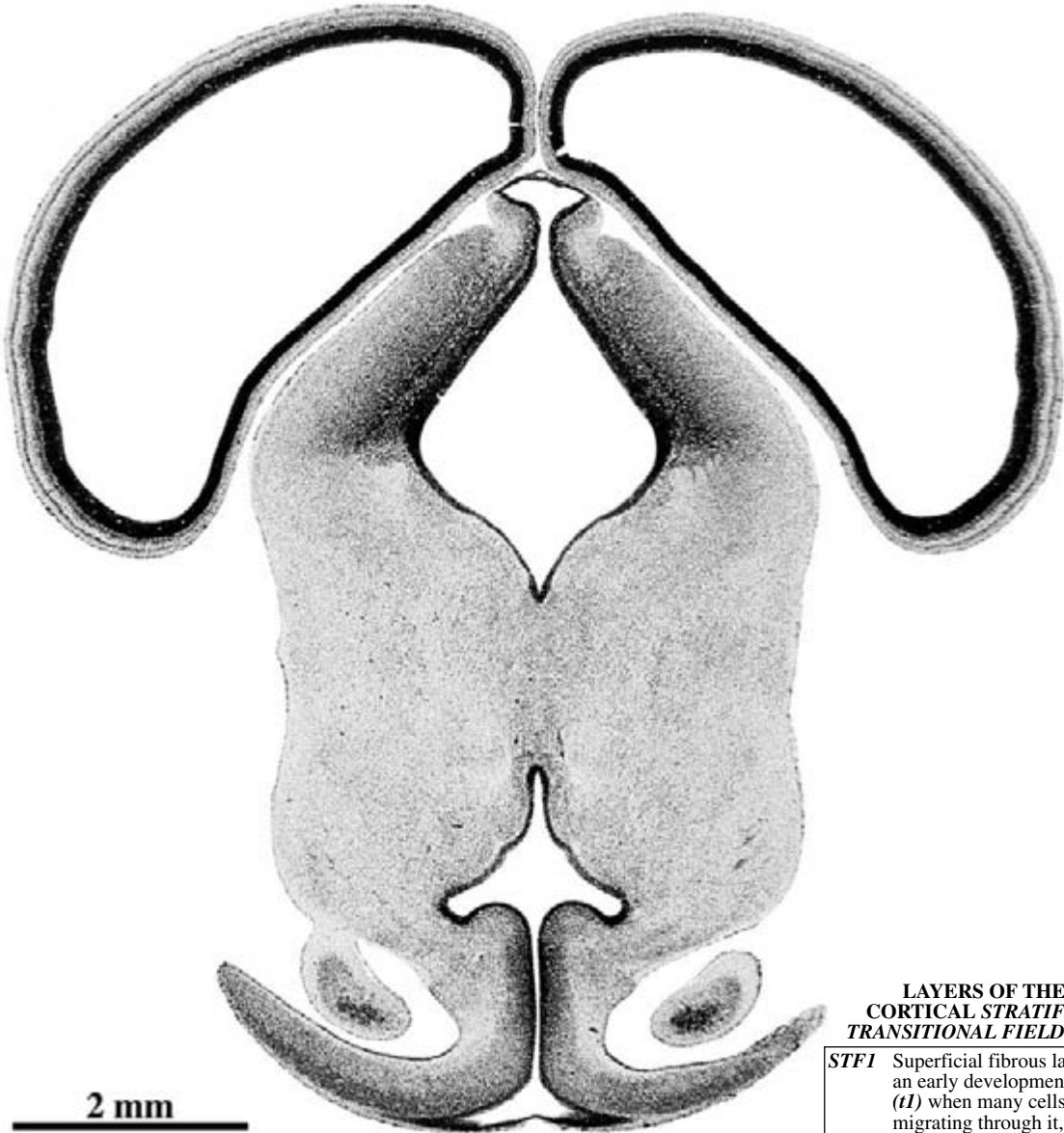
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glieoepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 163A

GW8 Coronal
CR 31 mm
C9226
Level 19:
Section 252

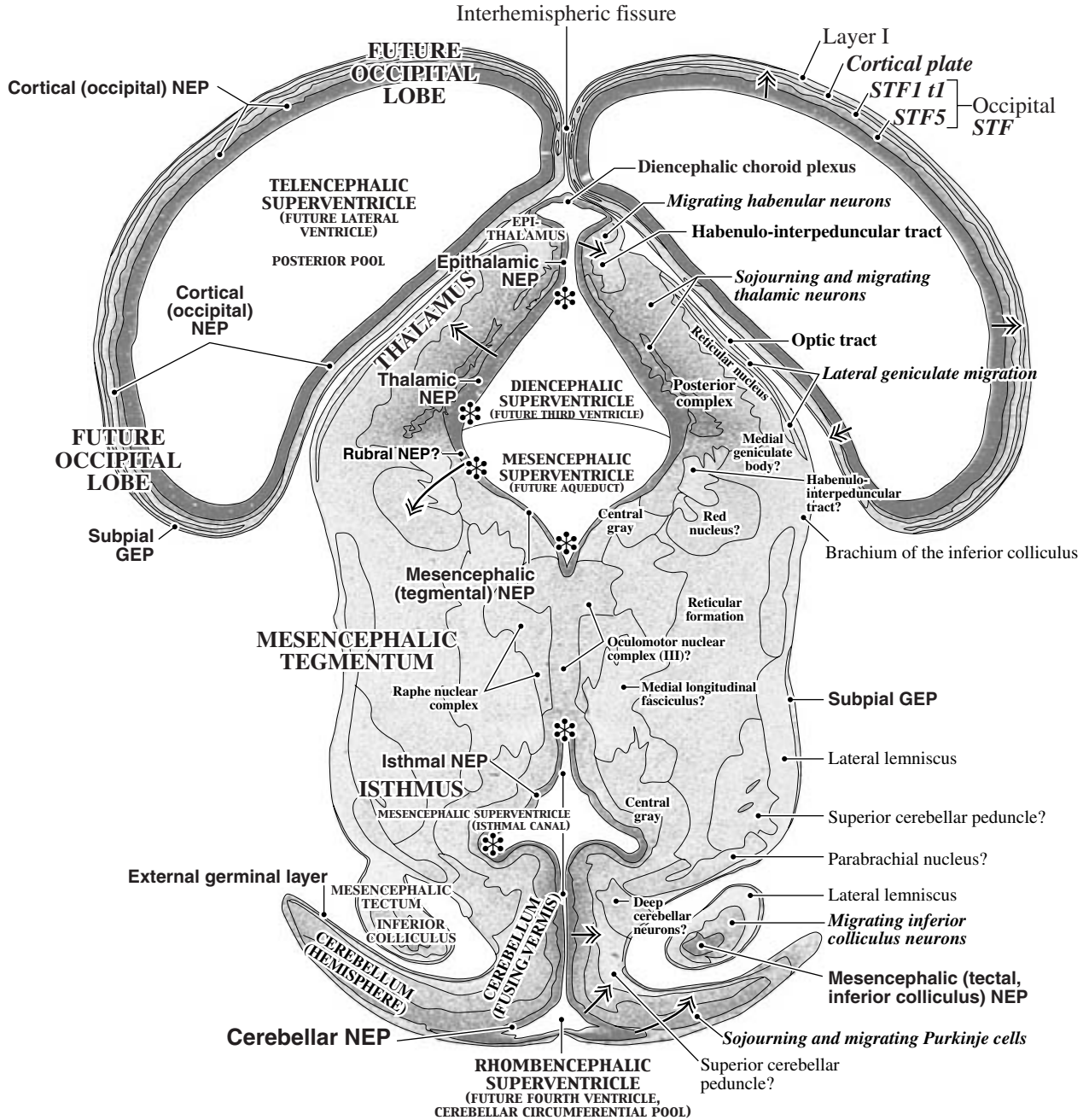


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 164A

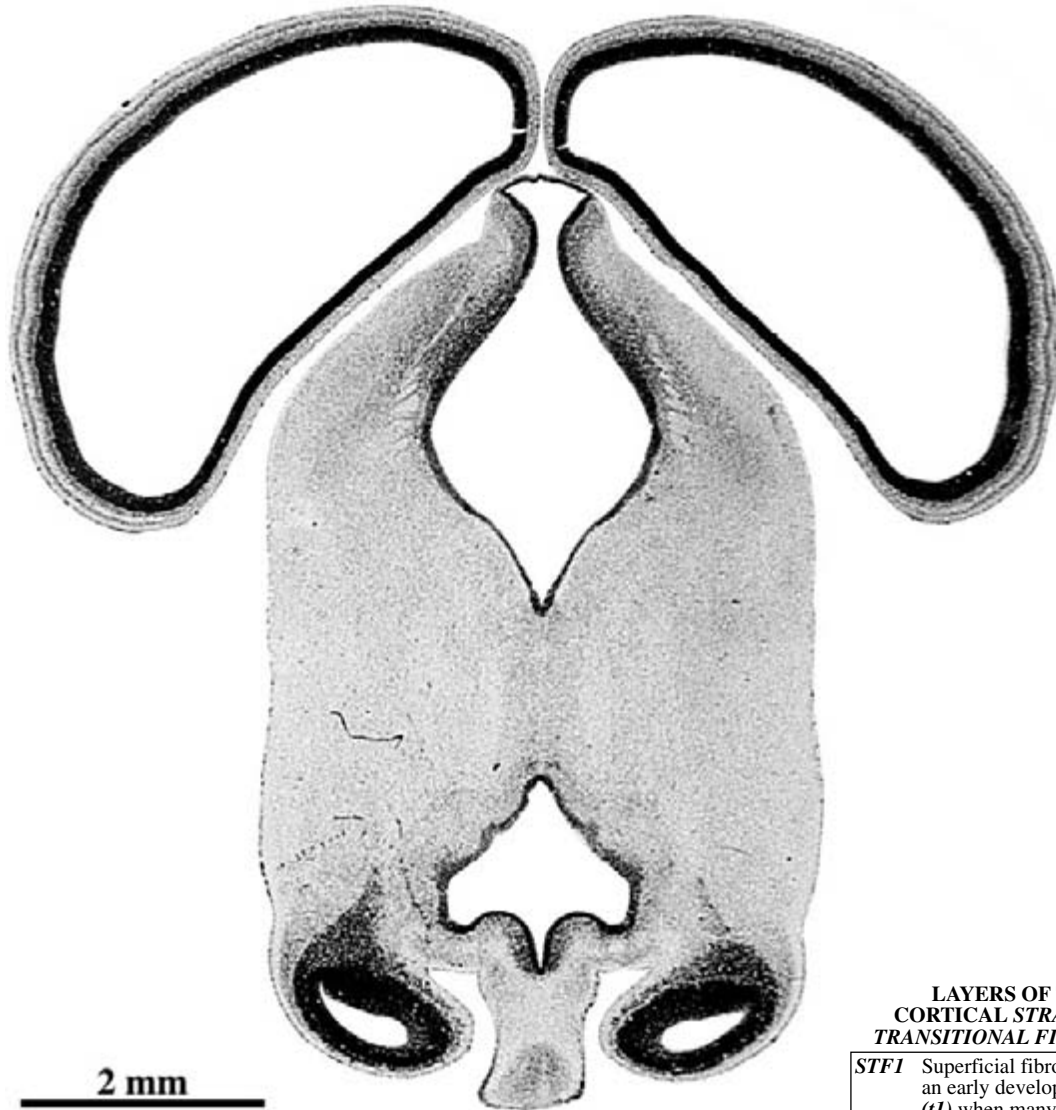
GW8 Coronal

CR 31 mm

C9226

Level 20:

Section 234

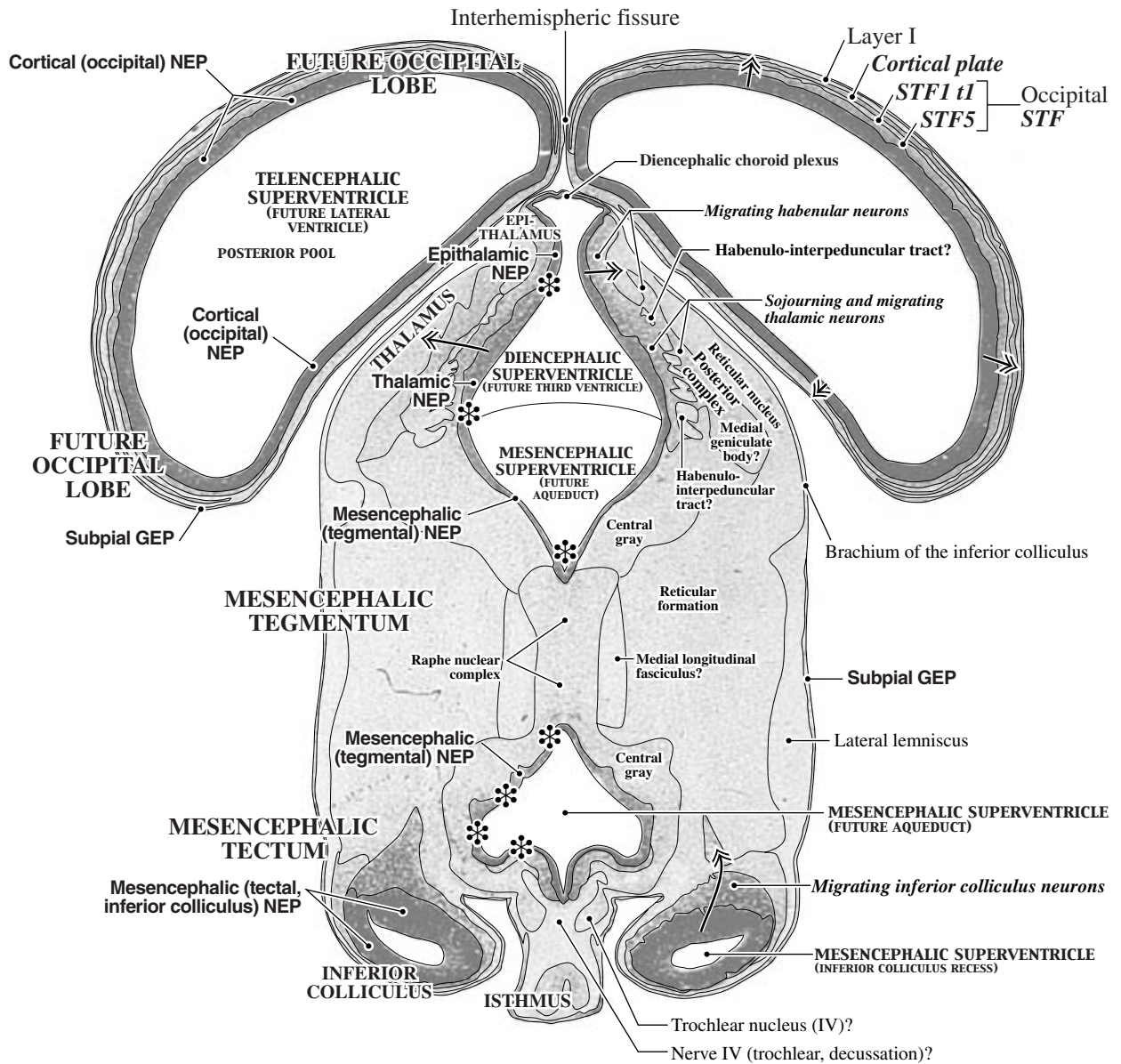


**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



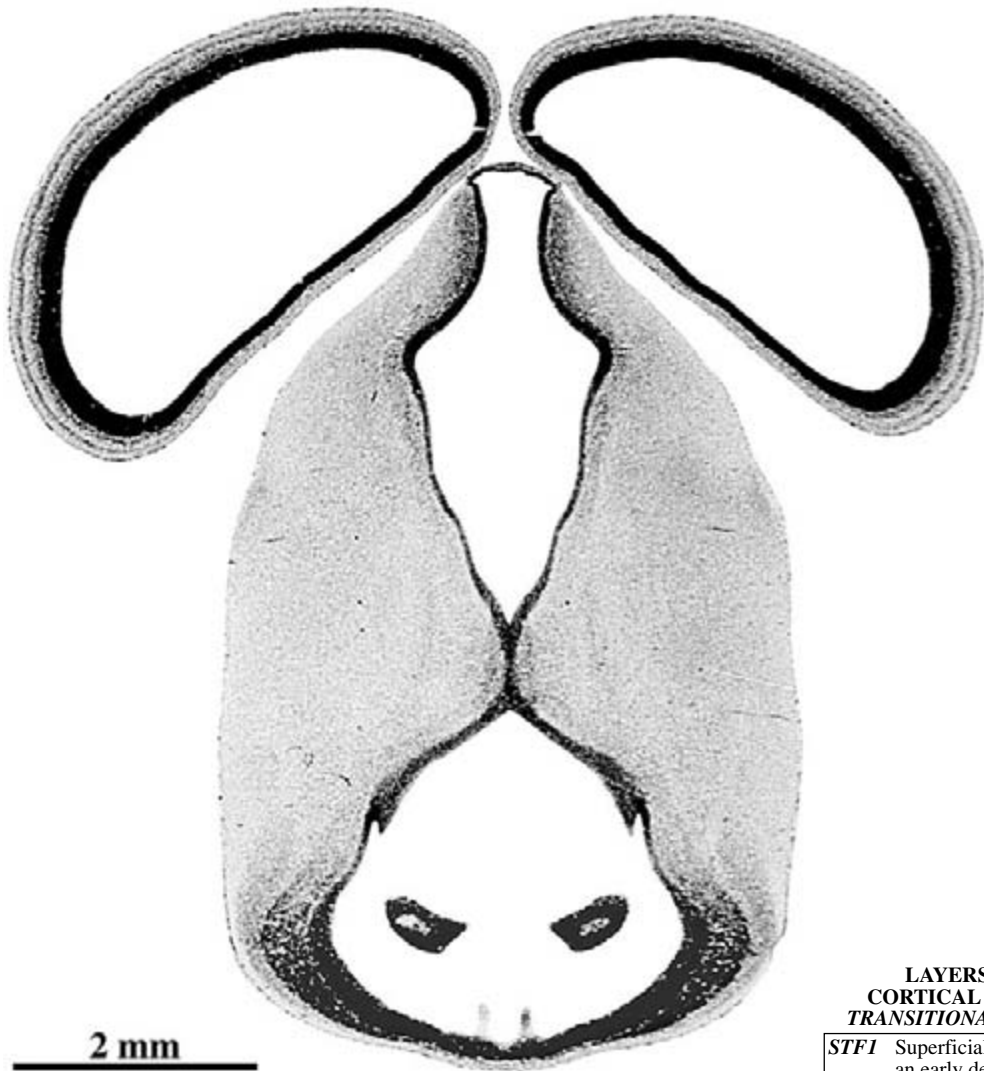
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 165A

GW8 Coronal
 CR 31 mm
 C9226
 Level 21:
 Section 206

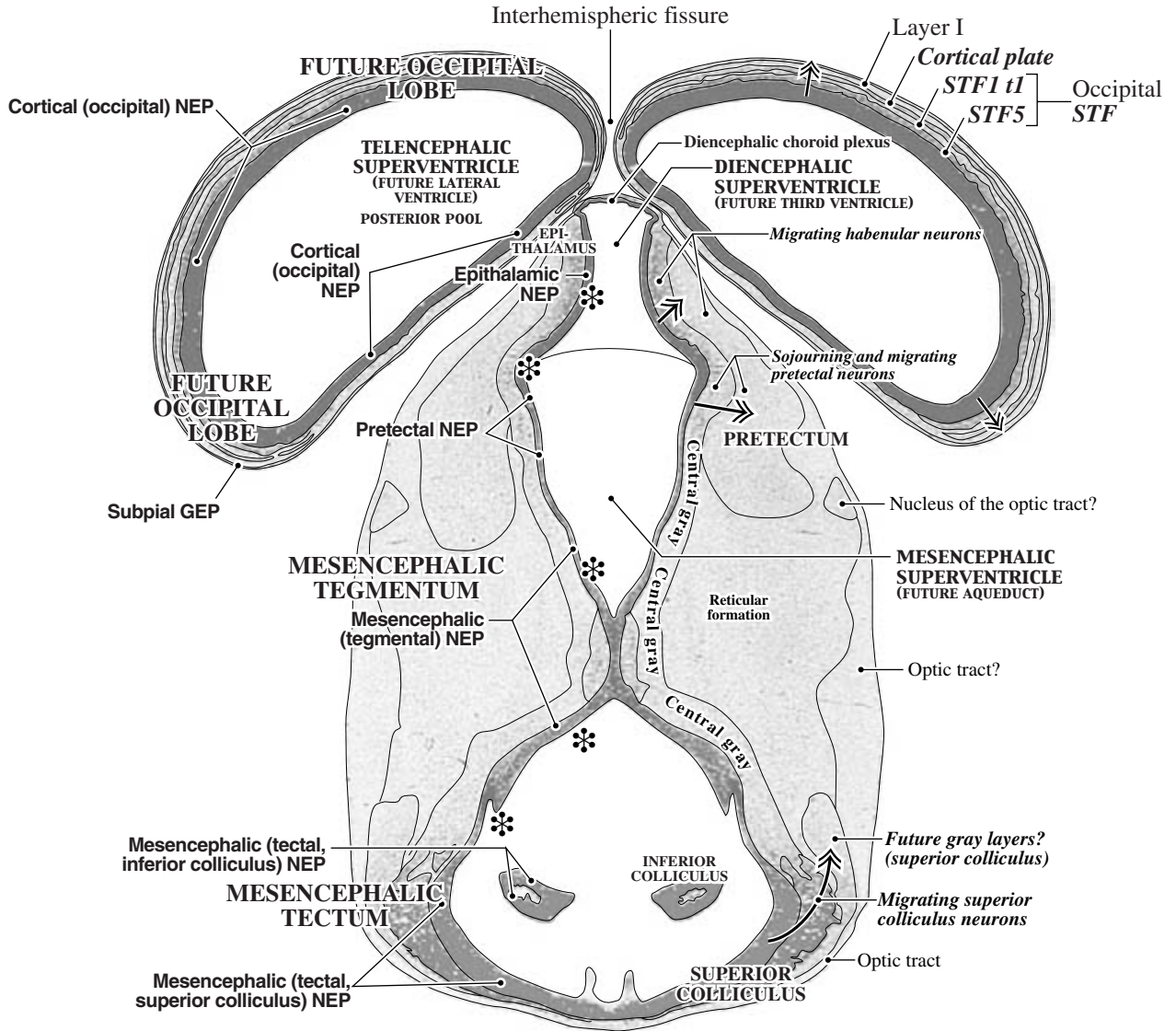


**LAYERS OF THE
 CORTICAL STRATIFIED
 TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**



* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 166A

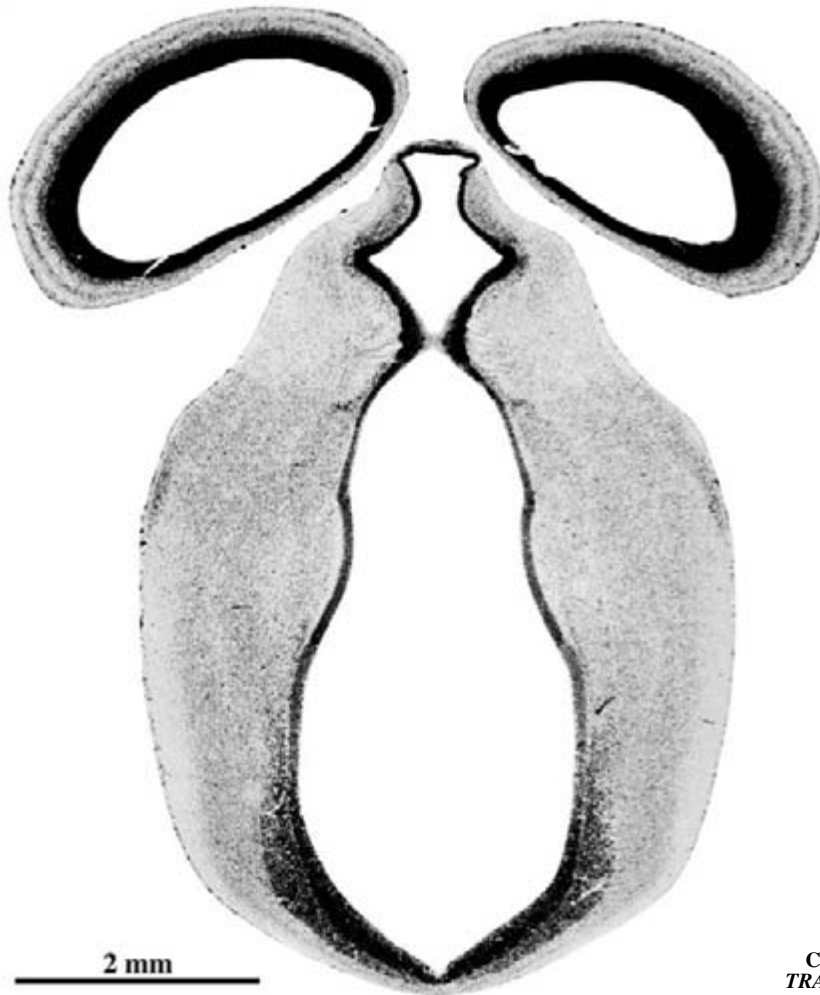
GW8 Coronal

CR 31 mm

C9226

Level 22:

Section 170



**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

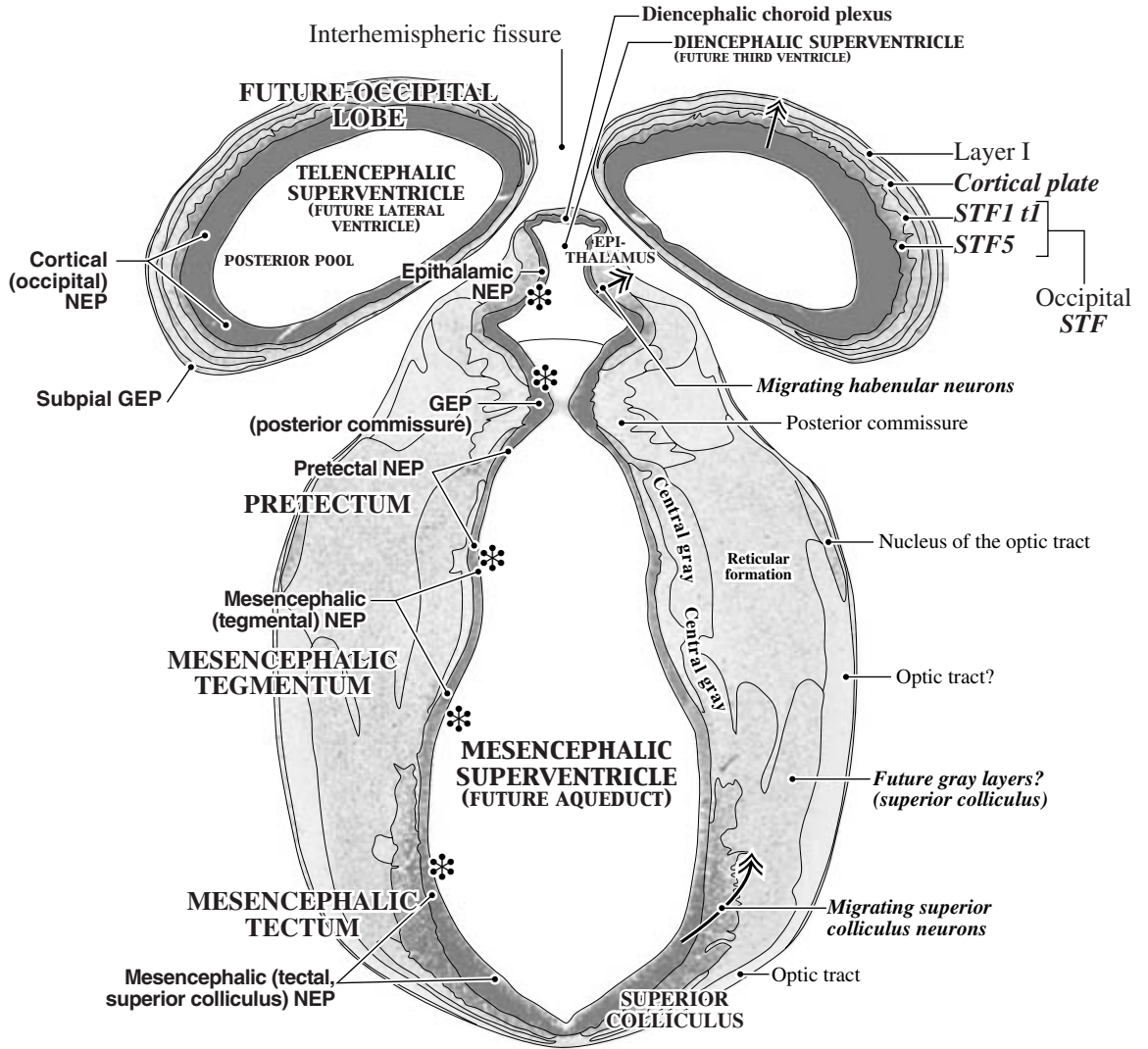
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or **Bold**



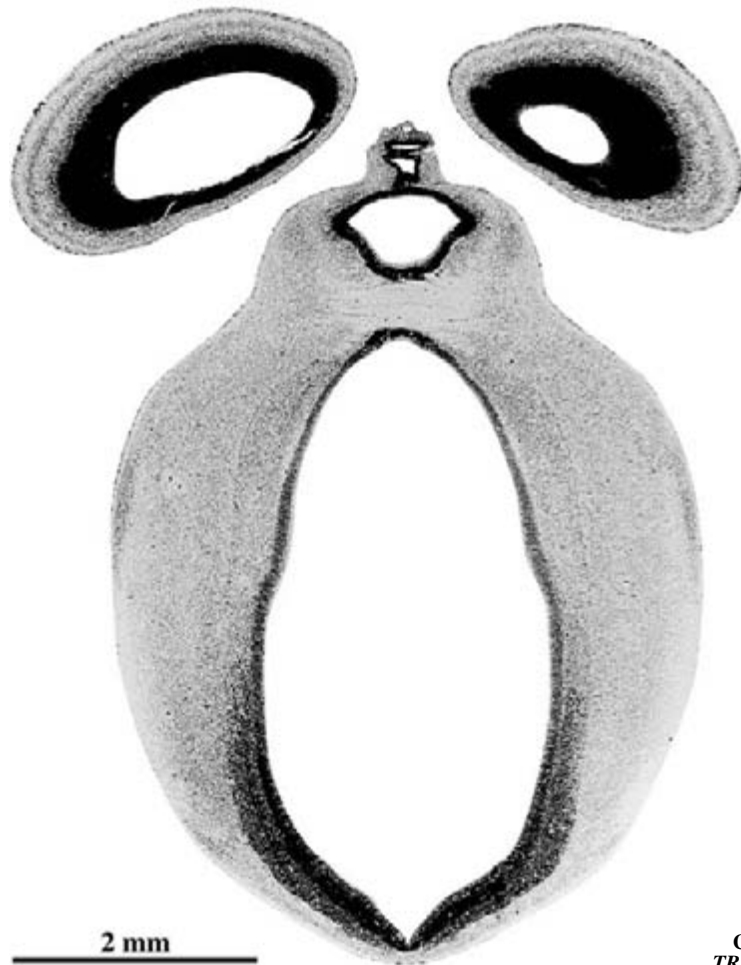
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 167A

GW8 Coronal
CR 31 mm
C9226
Level 23:
Section 158



**LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)**

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

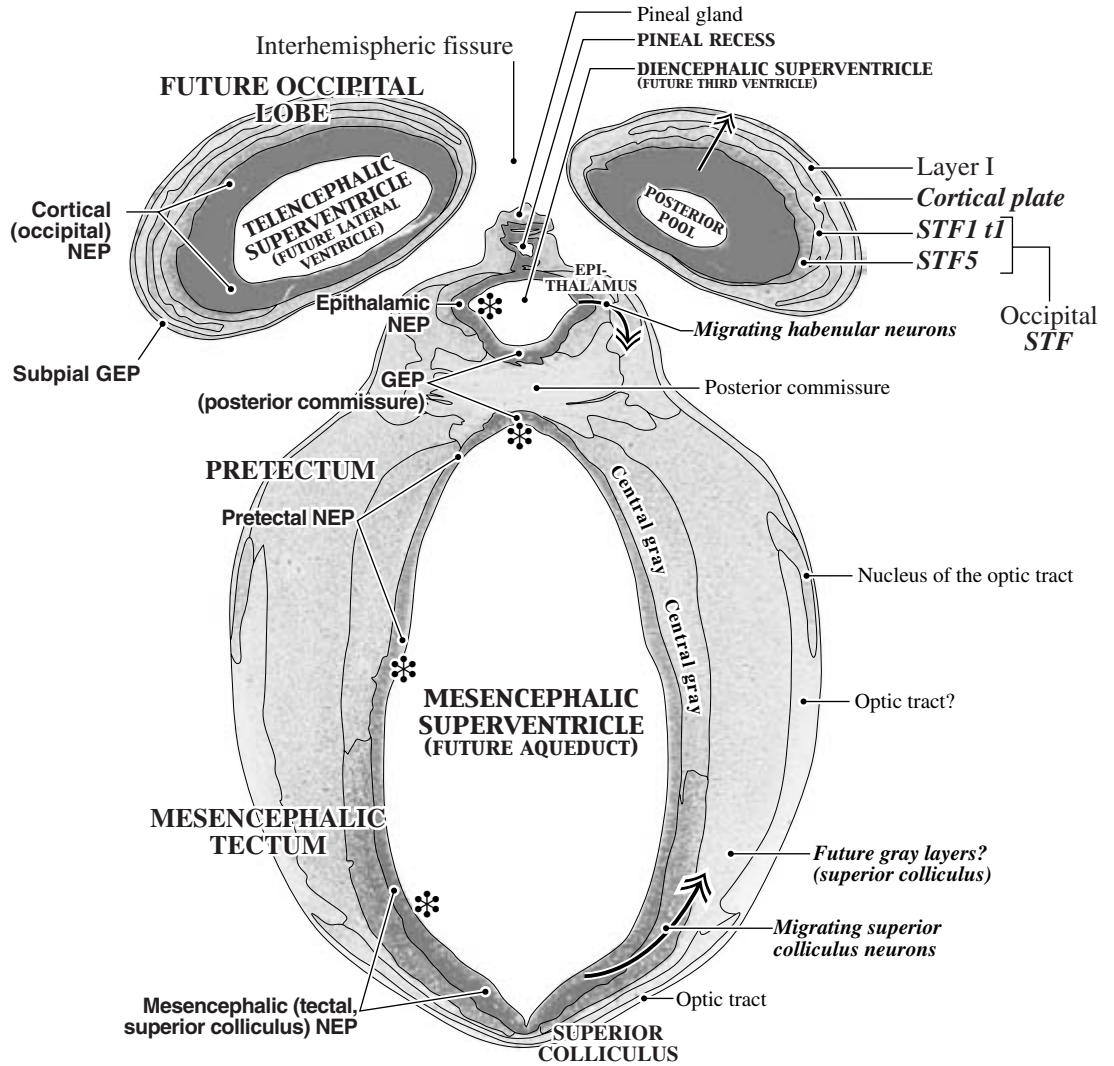
FONT KEY:

VENTRICULAR DIVISIONS - CAPITALS

Germinal zone - Helvetica bold

Transient structure - Times bold italic

Permanent structure - Times Roman or Bold



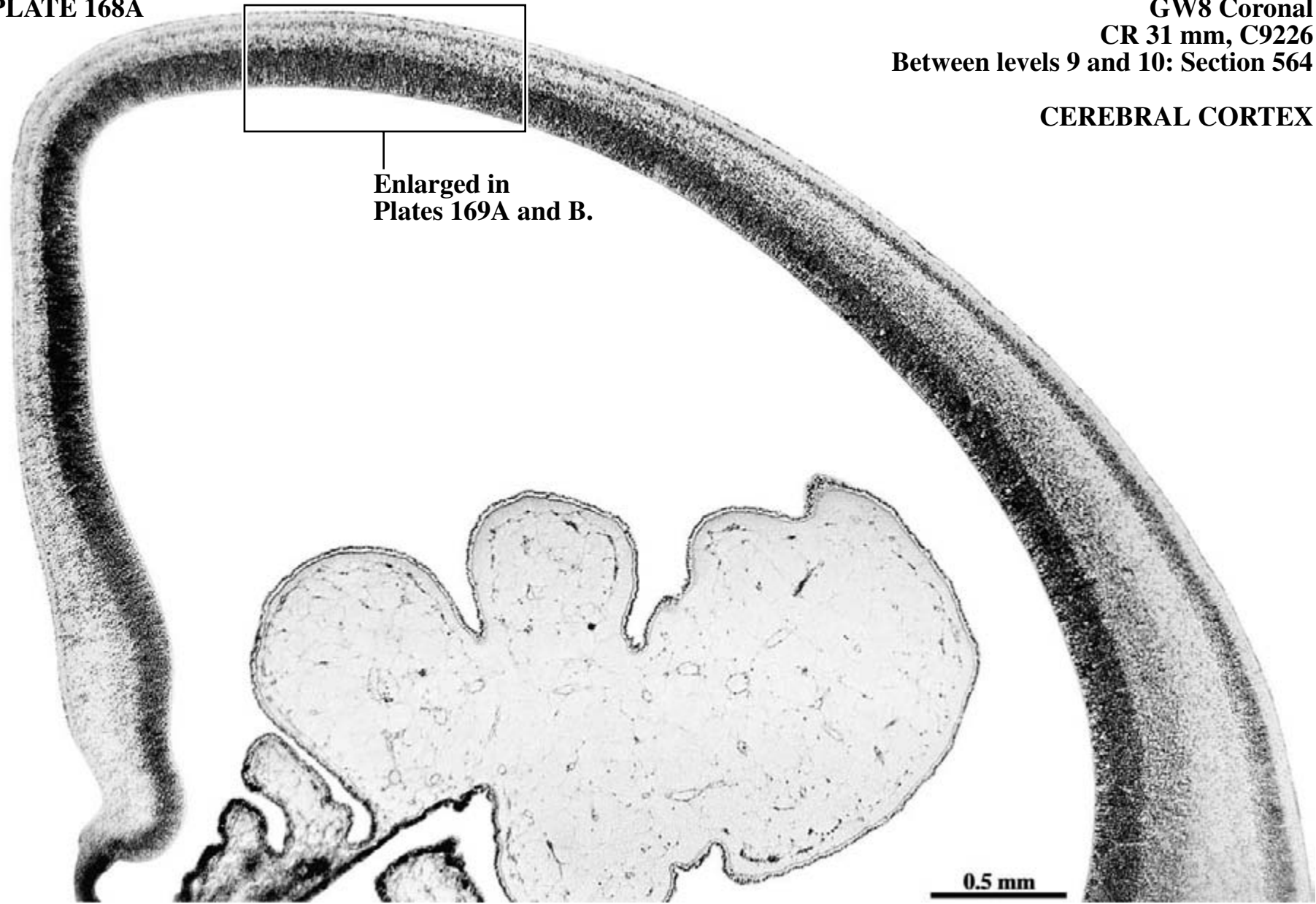
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

ABBREVIATIONS:
GEP - Glioeepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 168A

**GW8 Coronal
CR 31 mm, C9226
Between levels 9 and 10: Section 564
CEREBRAL CORTEX**



**Enlarged in
Plates 169A and B.**

0.5 mm

See levels 9 and 10 in Plates 153A and B to 154A and B.

PLATE 168B

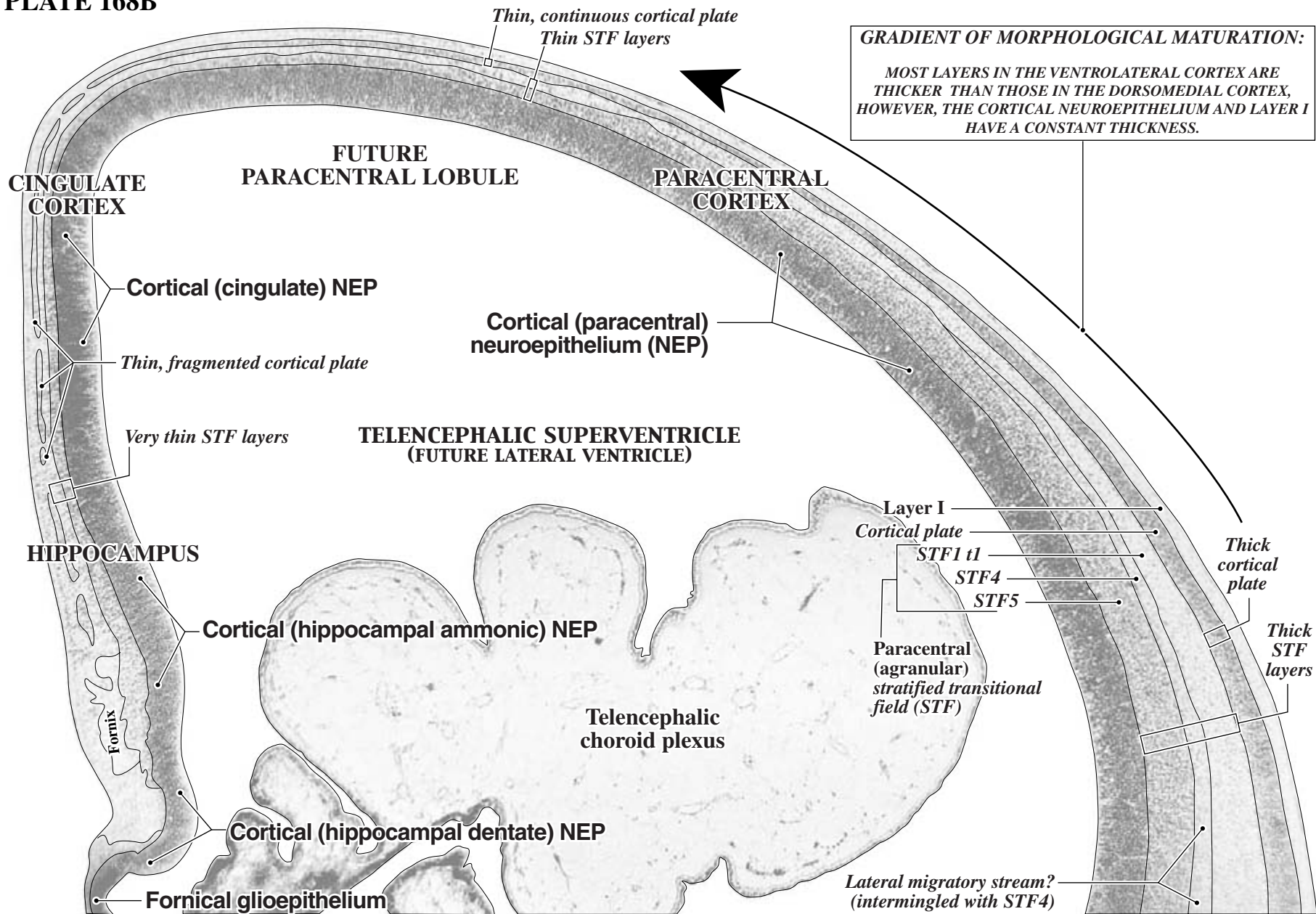
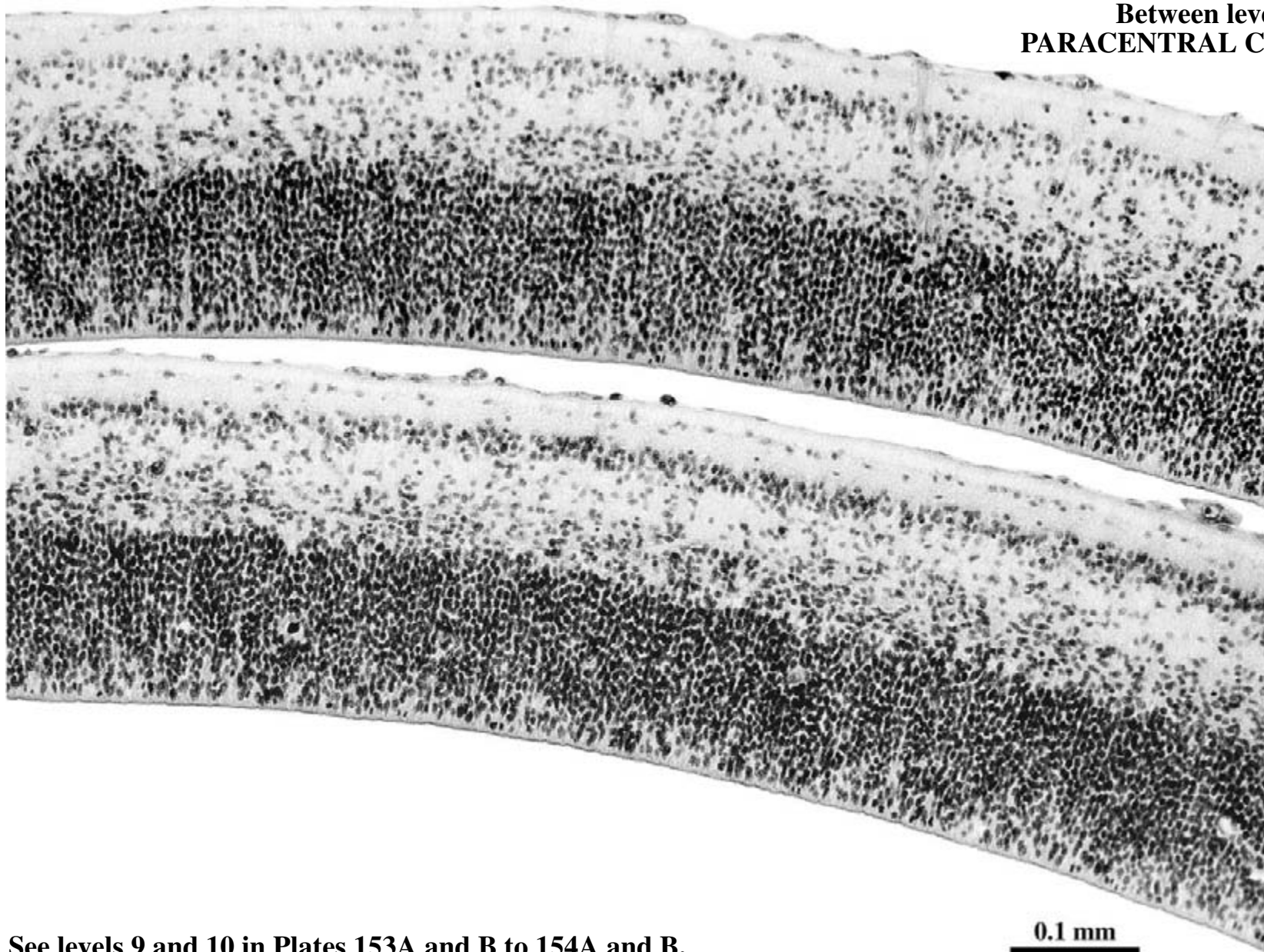


PLATE 169A

**GW8 Coronal, CR 31 mm, C9226
Between levels 9 and 10
PARACENTRAL CEREBRAL
CORTEX**



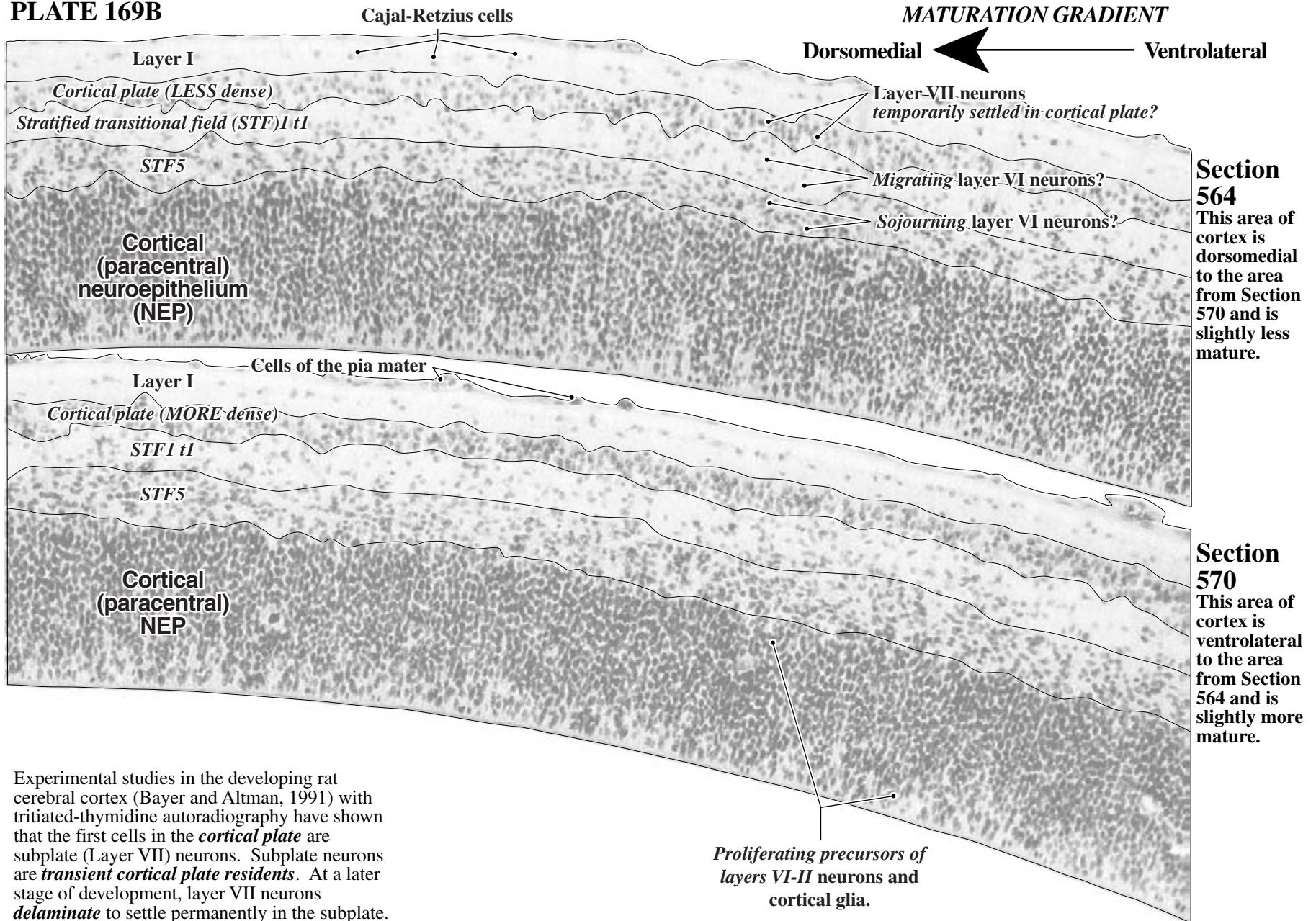
**Section
564**

**Section
570**

0.1 mm

See levels 9 and 10 in Plates 153A and B to 154A and B.

PLATE 169B



Experimental studies in the developing rat cerebral cortex (Bayer and Altman, 1991) with tritiated-thymidine autoradiography have shown that the first cells in the *cortical plate* are subplate (Layer VII) neurons. Subplate neurons are *transient cortical plate residents*. At a later stage of development, layer VII neurons *delaminate* to settle permanently in the subplate.

Section 564
This area of cortex is dorsomedial to the area from Section 570 and is slightly less mature.

Section 570
This area of cortex is ventrolateral to the area from Section 564 and is slightly more mature.

PART IX: GW8 HORIZONTAL

This is specimen number 609 in the Carnegie Collection, designated here as C609. A normal female fetus with a crown-rump length (CR) of 32 mm was collected in 1916, and is estimated to be in gestational week (GW) 8. The entire fetus was embedded in paraffin, cut transversely in 50 μ m thick sections, and stained with aluminum cochineal. Since there is no photograph of this brain before it was embedded and cut, a specimen from Hochstetter (1919) that is only partially comparable to C609 has been modified to show the approximate section plane and external features of the brain at GW8 (**Figure 8**). Like most of the specimens in this Volume, the sections are not cut exactly in one plane; C609's cortex is cut midway between coronal and horizontal planes, and is presented as a "horizontal" brain. The C609 section planes through the cortex and brainstem are not at the same angle when transferred to Hochstetter's CR27 mm specimen. Instead, brainstem planes of section appear to fan upward and downward from sections in the cortex apparently around a fulcrum centering in the invagination of the medullary velum overlying the rhombencephalic superventricle. We interpret this to indicate that the brain flexures are more loosely folded in the Hochstetter specimen than in C609. But it is difficult to determine how the brainstem is folded in C609 to make the section planes line up with those in the cortex. Photographs of 23 sections (**Levels 1-10**) are illustrated at low magnification in **Plates 170-179**. High-magnification views of different areas of the brain are shown in **Plates 180-185**. To maximize image size within page space, all of C609's sections are rotated 90° (landscape orientation). The anterior part of each section is on the left (page bottom), and the posterior part of each section is on the right (page top).

C609 is similar to the other GW8 specimens and shows brain maturation in still another perspective. The telencephalic and rhombencephalic *superventricles* are obvious, along with the slit-like diencephalic and mesencephalic superventricles. The parenchyma, the area between the superficial border of the *neuroepithelium* (NEP) / *subventricular zone* (SVZ) and the pial membrane, is the region where neurons migrate, settle, and differentiate. The thicknesses of the neuroepithelium and the parenchyma are similar to those in C9226 throughout the brain indicating brain maturation in both specimens is similar.

The parenchyma is thick and bordered by a thin NEP in the medulla, pons, and midbrain tegmentum without sur-

rounding dense sojourn zones. Most neurons have been generated here, few are migrating, and most are settled and differentiating. The two exceptions seen in C9226 are also seen in C145. First, presumptive facial motor neurons are clumped near the pontomedullary trench and some are migrating toward their ventral pontine/medullary settling sites. Second, the thicker *precerebellar neuroepithelium* in the medulla is generating predominantly pontine gray neurons; many precerebellar neurons are migrating in the *anterior and posterior extramural migratory streams*. The cerebellar NEP is thicker than that in the pons and medulla, and the cerebellar parenchyma has a dense Purkinje cell sojourn zone below presumptive earlier-generated deep neurons; the *external germinal layer* (*egl*) is rudimentary. The mesencephalic tectal NEP is thick adjacent to a thin parenchyma that contains dense sojourning and migrating neurons; substantial neurogenesis is ongoing in both the superior and inferior colliculi in the midbrain tectum.

The prominent diencephalic NEP and thick parenchyma filled with dense zones of sojourning and migrating neurons is remarkable in C609. Many migratory streams are visible in the thalamus, and migrating subthalamic nucleus neurons can be followed from the posterior hypothalamic NEP to the subthalamic nucleus. Although many diencephalic neurons have been generated by GW8, most of them are still migrating and few have settled. In spite of that, there are large accumulations of fibers (presumably thalamic axons) in the lateral thalamus and internal capsule, indicating that young thalamic neurons grow axons toward the cerebral cortex before settling and differentiating.

Within the telencephalon, the cerebral cortex has a thick NEP and a very thin parenchyma, indicating that most of its neurons are still not generated. The cerebral cortical NEP is the sole germinal matrix. The *stratified transitional field* (*STF*) contains *STF1* and *STF5* only in lateral areas. The pronounced anterolateral (thicker) to dorsomedial (thinner) maturation gradient is evident in the cerebral cortex. The basal telencephalic NEP/SVZ and parenchyma are both thick because there are large early-generated neuronal populations (for example, globus pallidus and substantia innominata) and massive late-generated populations (striatal neurons in the caudate and putamen). Most of the neurons settling in the basal telencephalon at GW8 are those of the early-generated populations, while many striatal neurons have not been generated yet.

GW8 "HORIZONTAL" SECTION PLANES

C609's cutting angle in the cerebral cortex rotates 45° counterclockwise from the true coronal plane (90°), exactly between true horizontal and true coronal.

This brain is less mature (CR 27 mm) with more loosely folded flexures than C609's brain (CR 32 mm). That is why cutting planes in C609's brainstem differ considerably from those in the cortex, many are horizontal. In the illustrated sections on the following pages, the anterior part of each section (left side) is dorsal to the posterior part (right side).

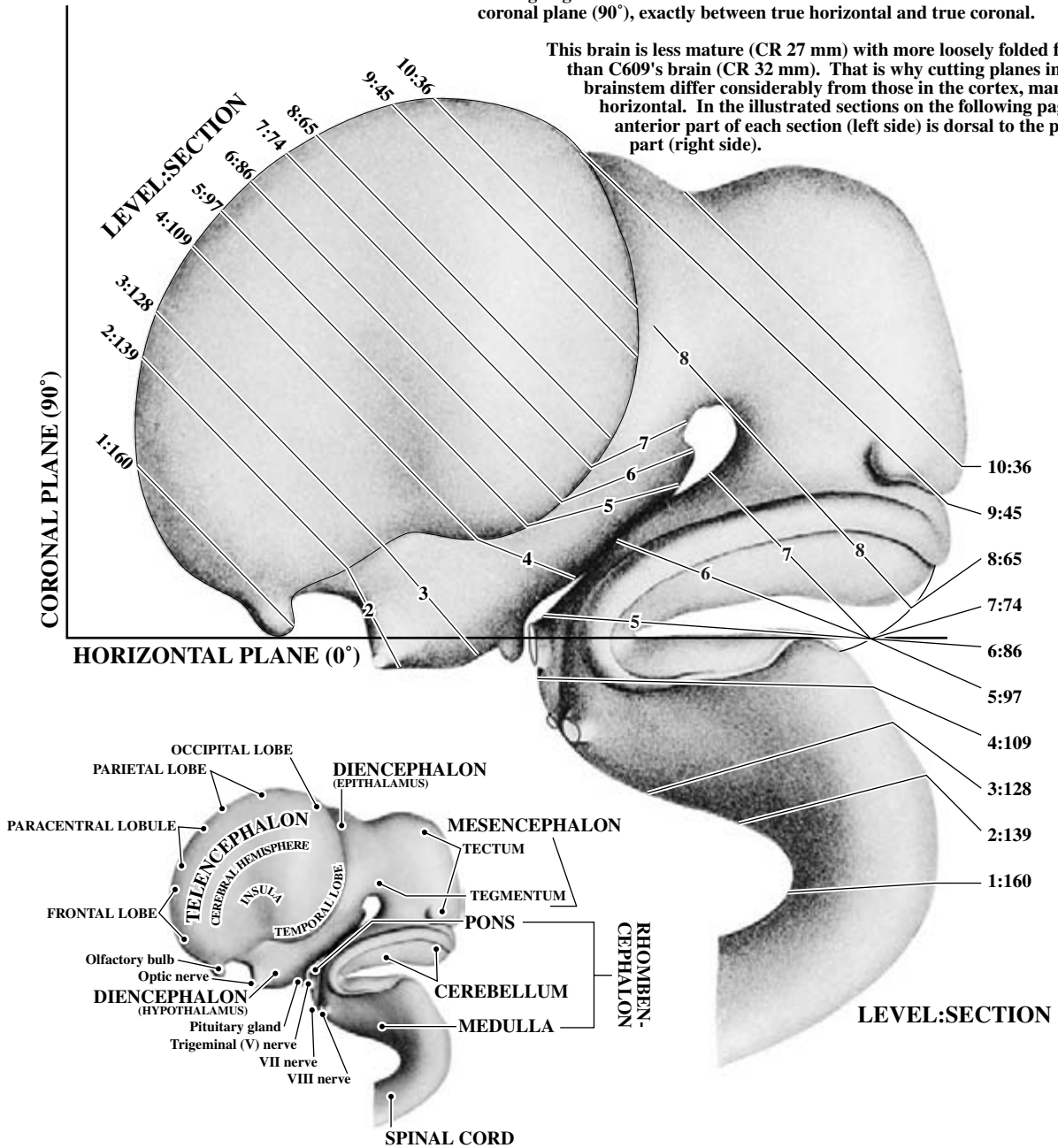


Figure 8. The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 27 mm (modified from Figure 37, Table VII, Hochstetter, 1919) serves to show the approximate locations and cutting angles of the illustrated sections of C609 in the following pages. The small inset identifies the major structural features. The line in the cerebellum and dorsal edges of the pons and medulla is the cut edge of the medullary velum.

PLATE 170A

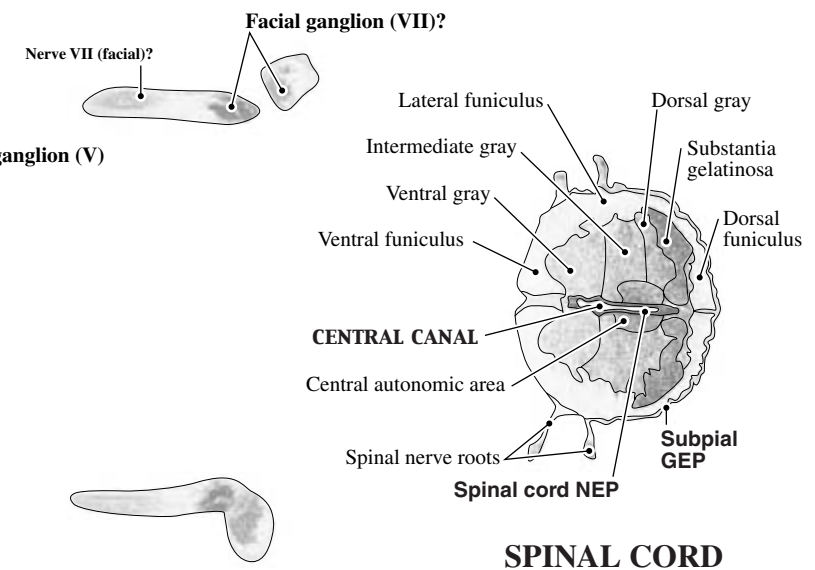
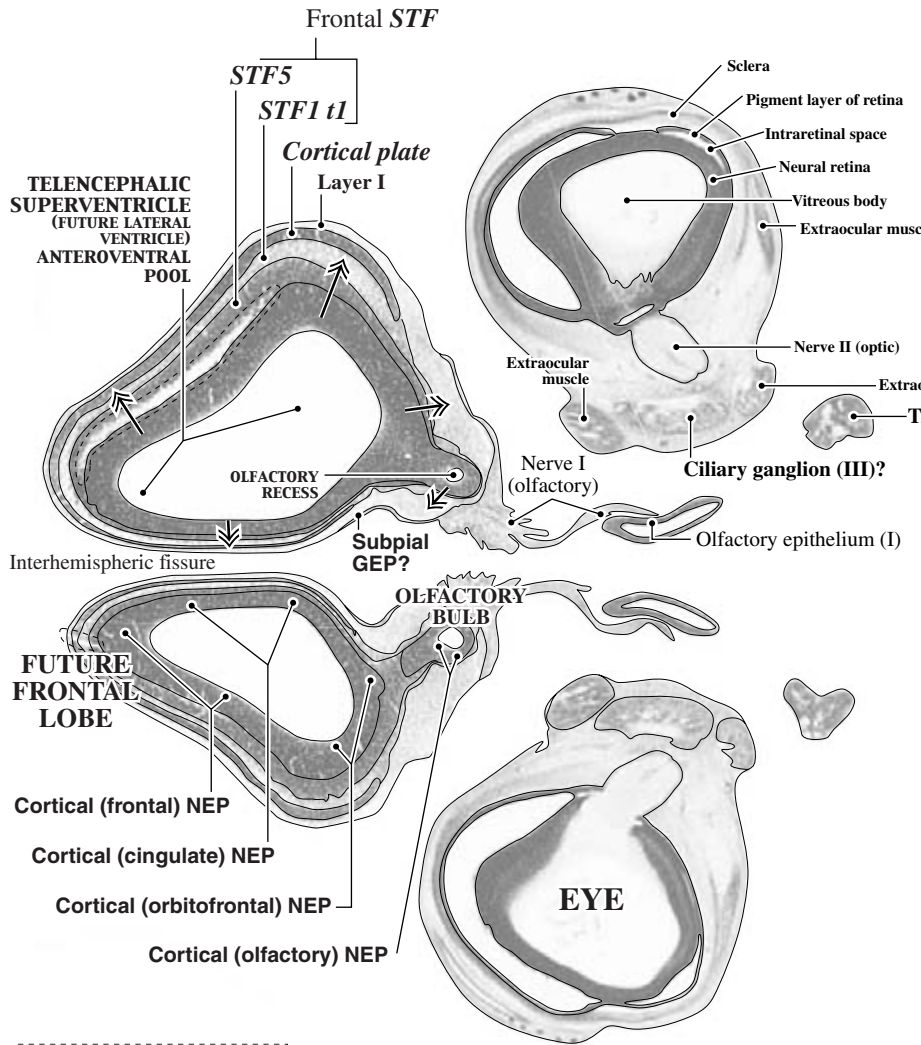
**GW8 Horizontal
CR 32 mm
C609
Level 1:
Section 160**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 170B



Dashed lines indicate staining and/or sectioning artifacts.

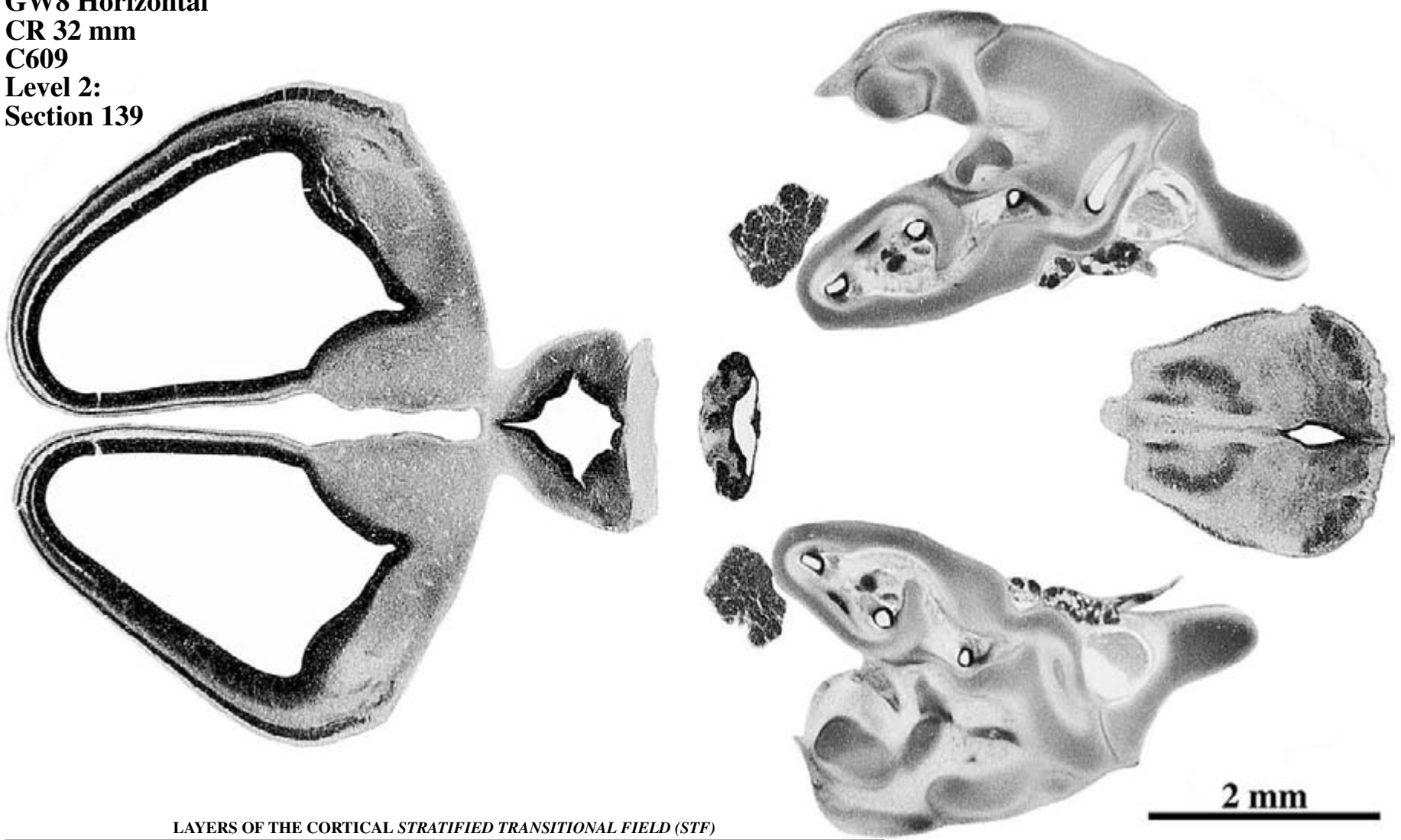
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 171A

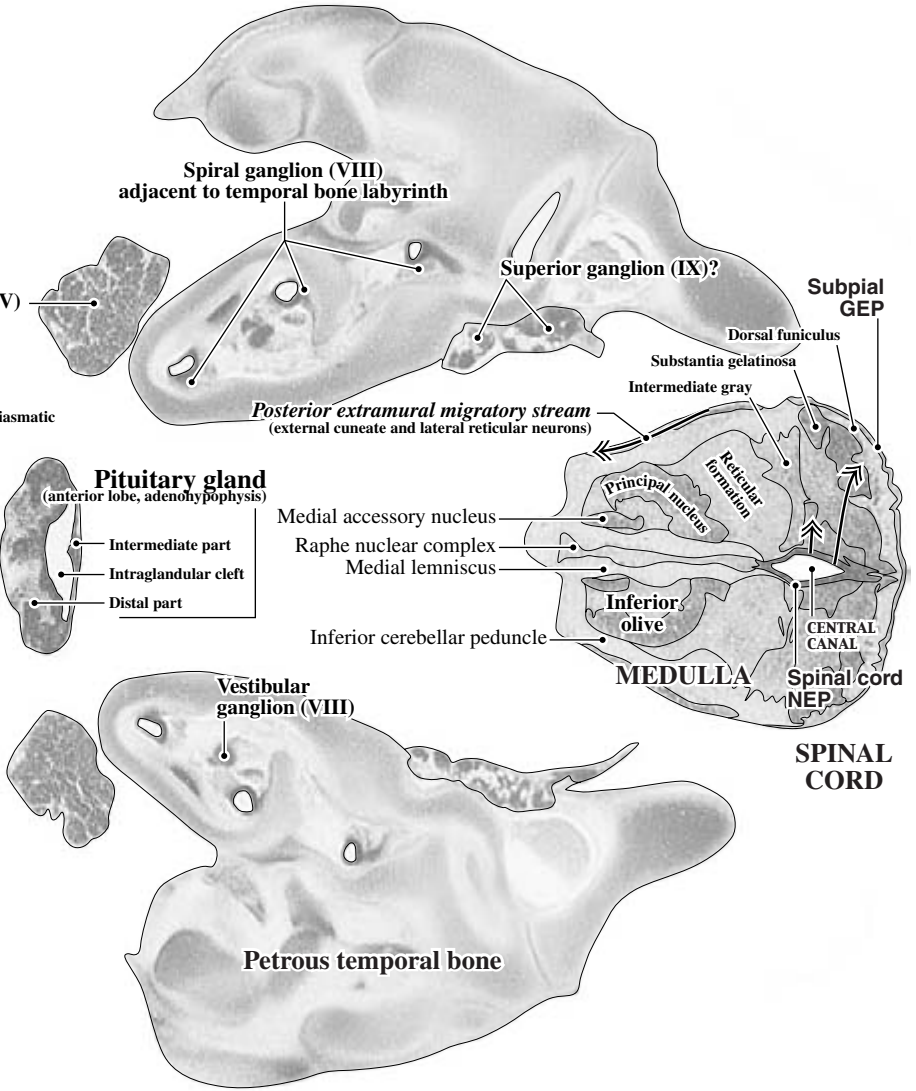
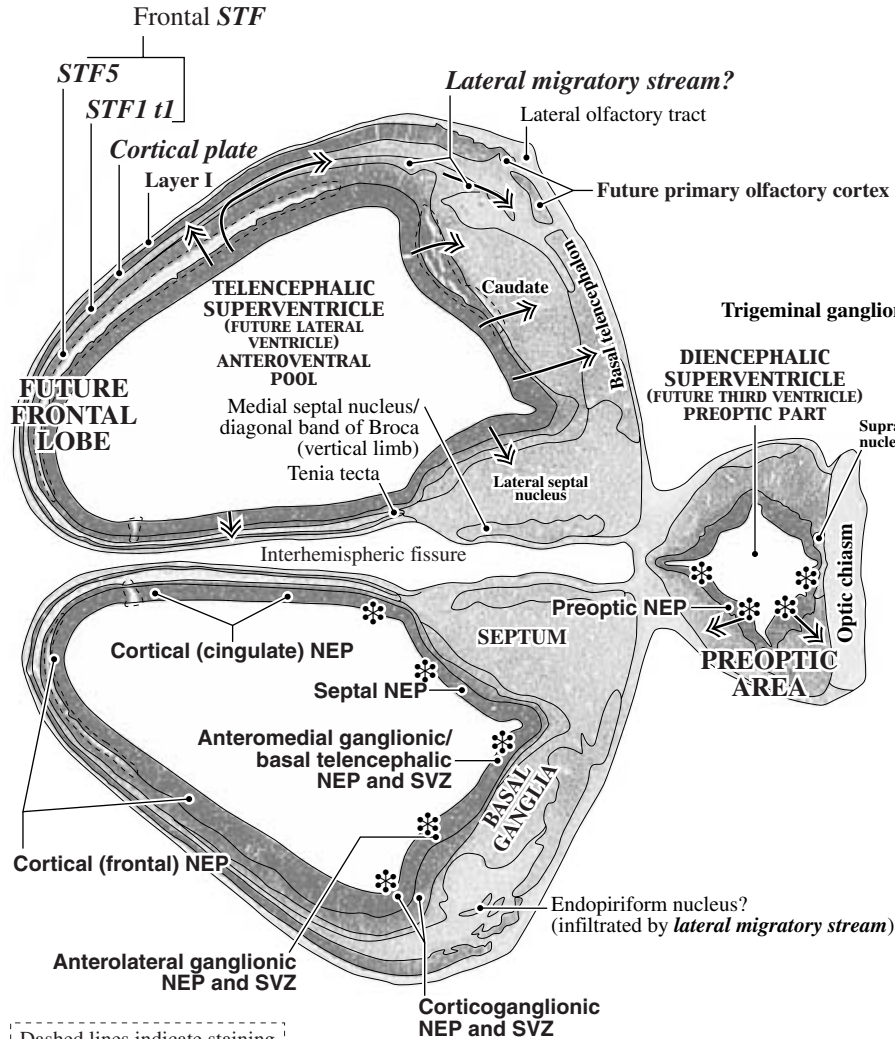
**GW8 Horizontal
CR 32 mm
C609
Level 2:
Section 139**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 171B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

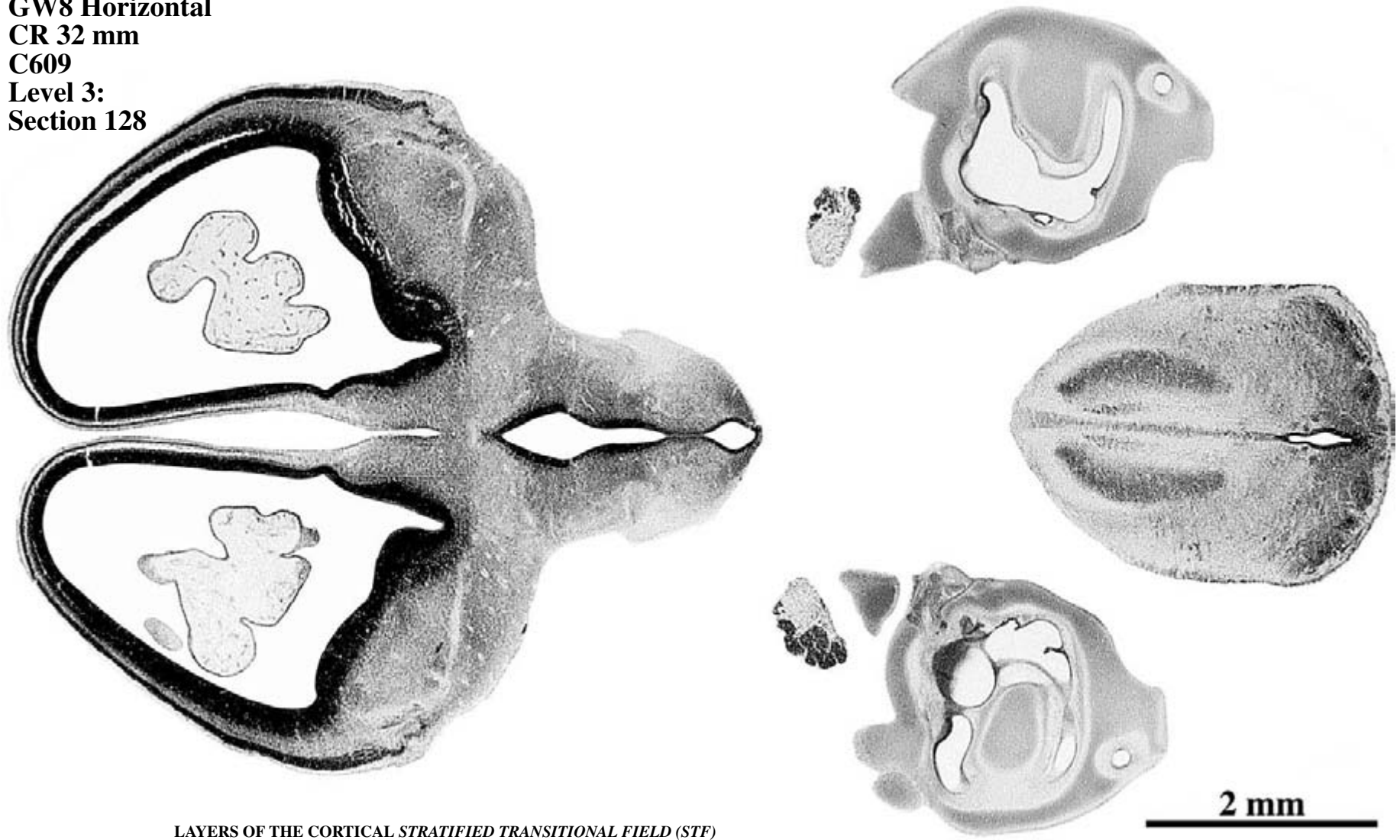
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 172A

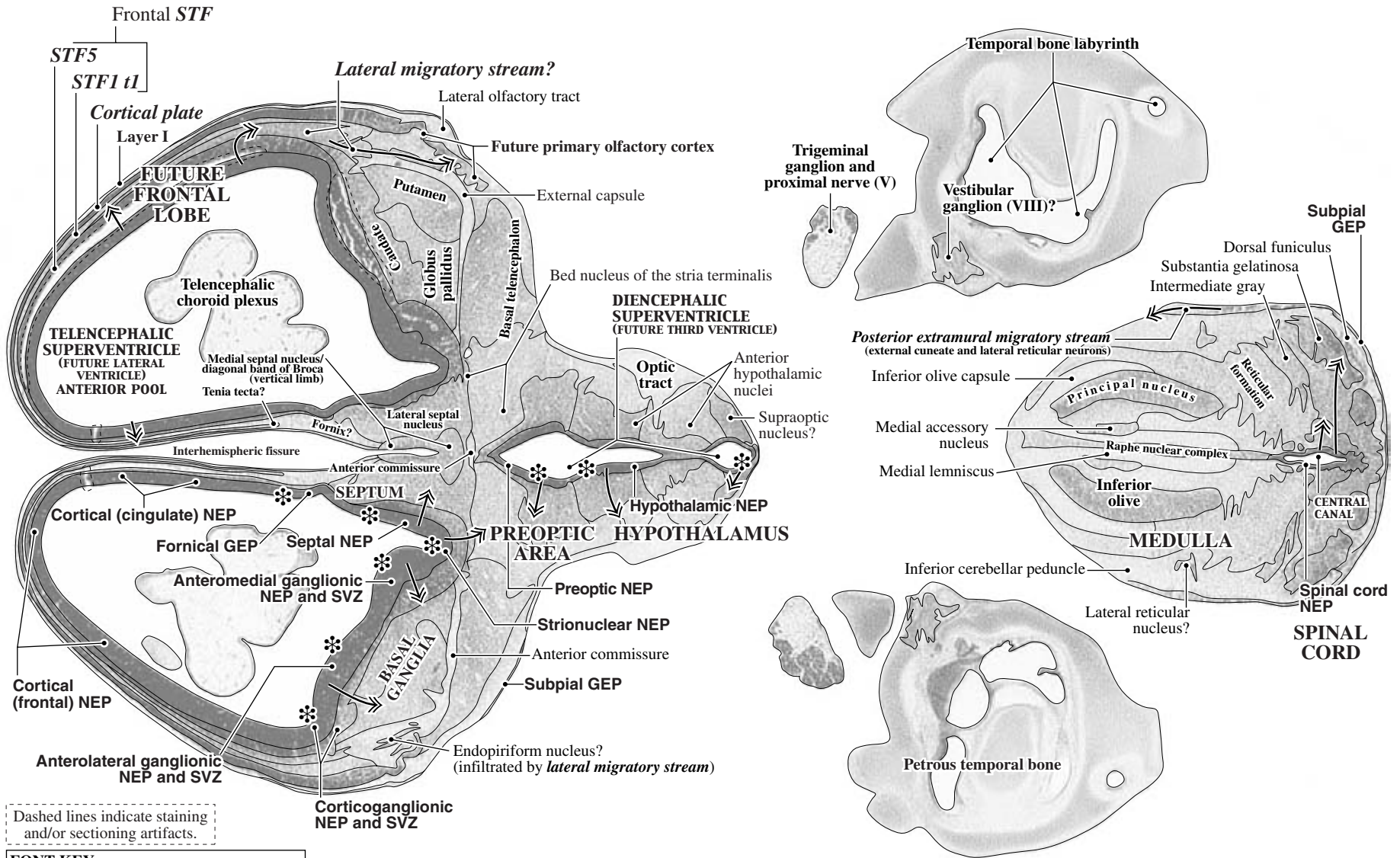
**GW8 Horizontal
CR 32 mm
C609
Level 3:
Section 128**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 172B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

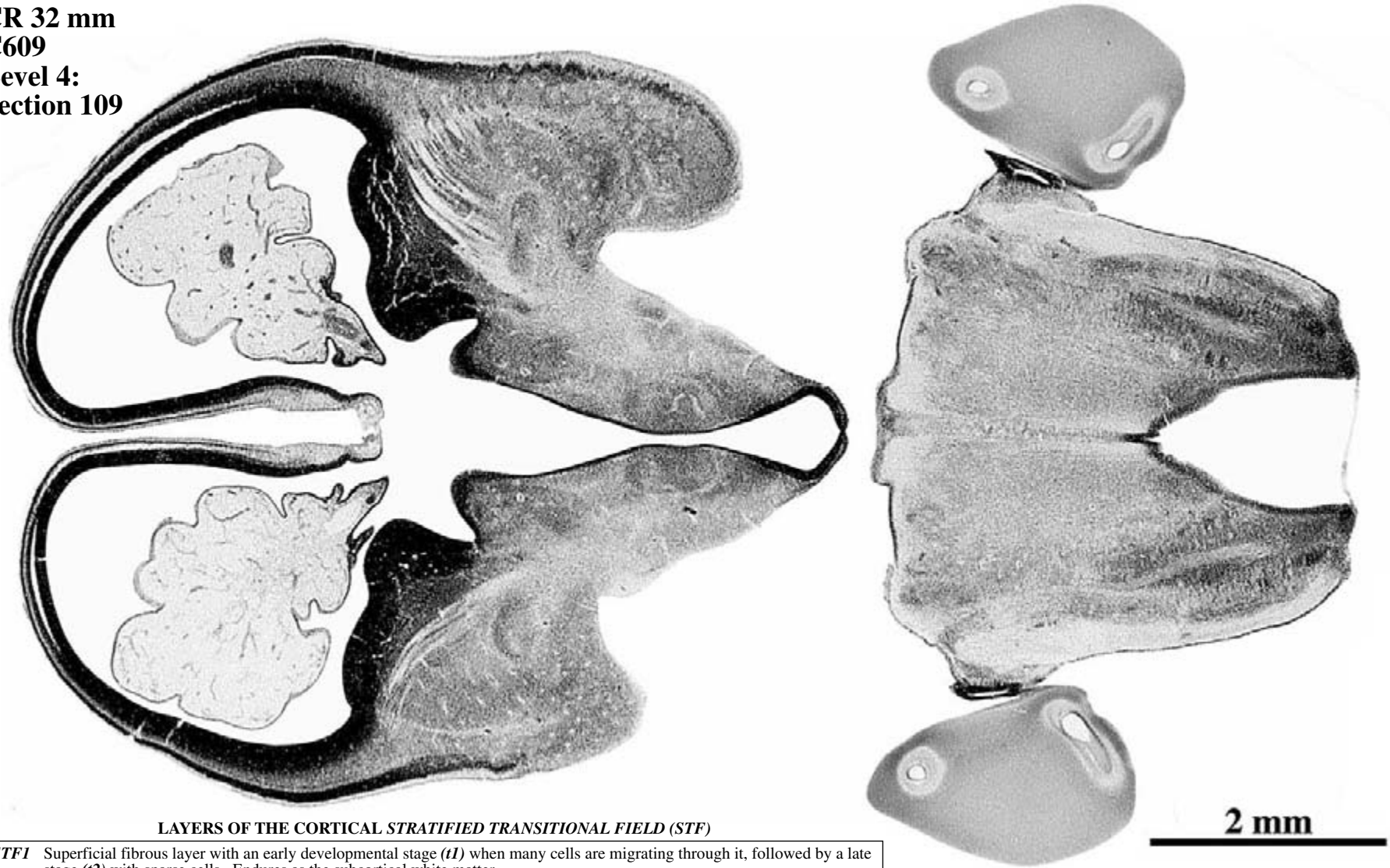
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 173A

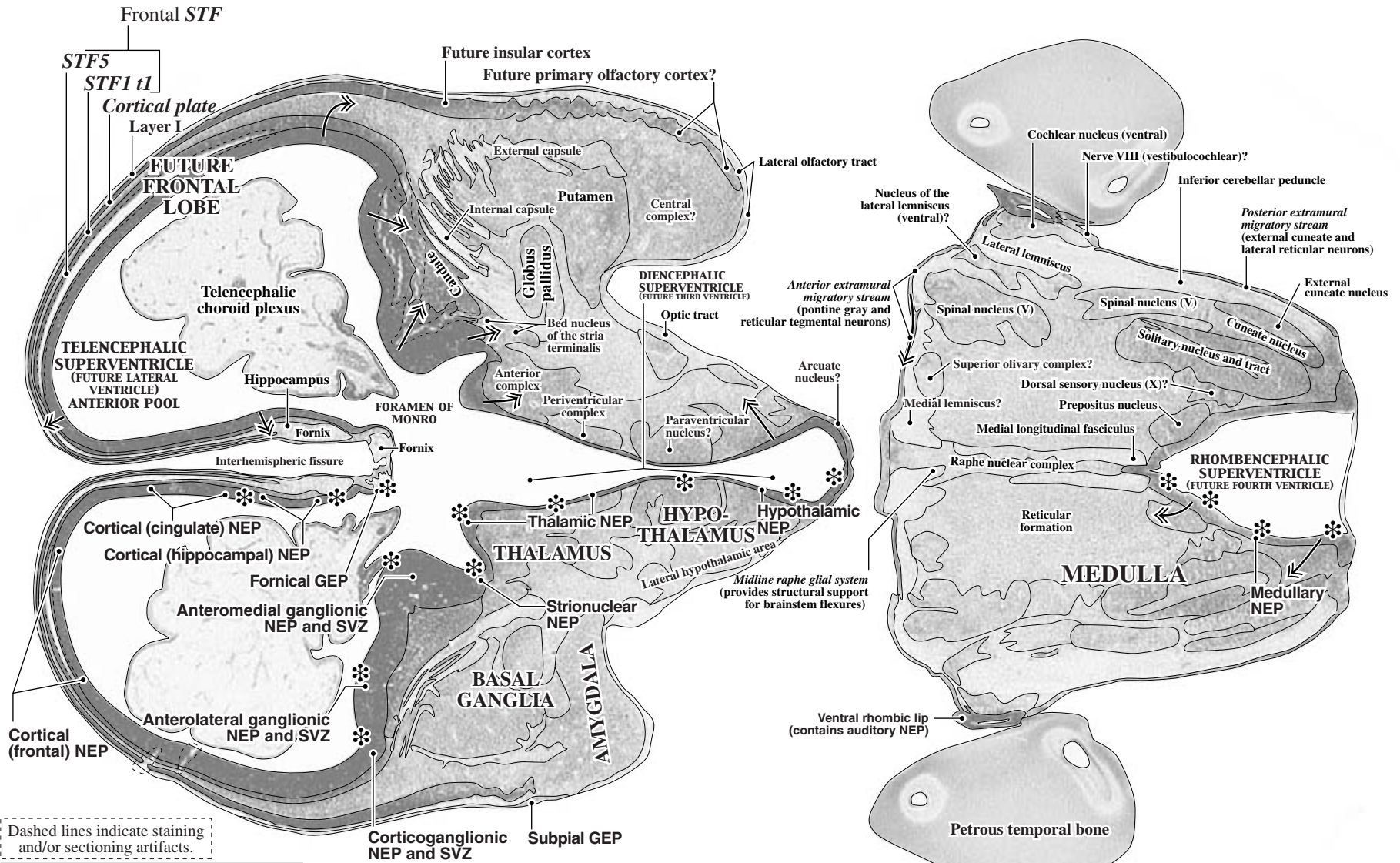
**GW8 Horizontal
CR 32 mm
C609
Level 4:
Section 109**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 173B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

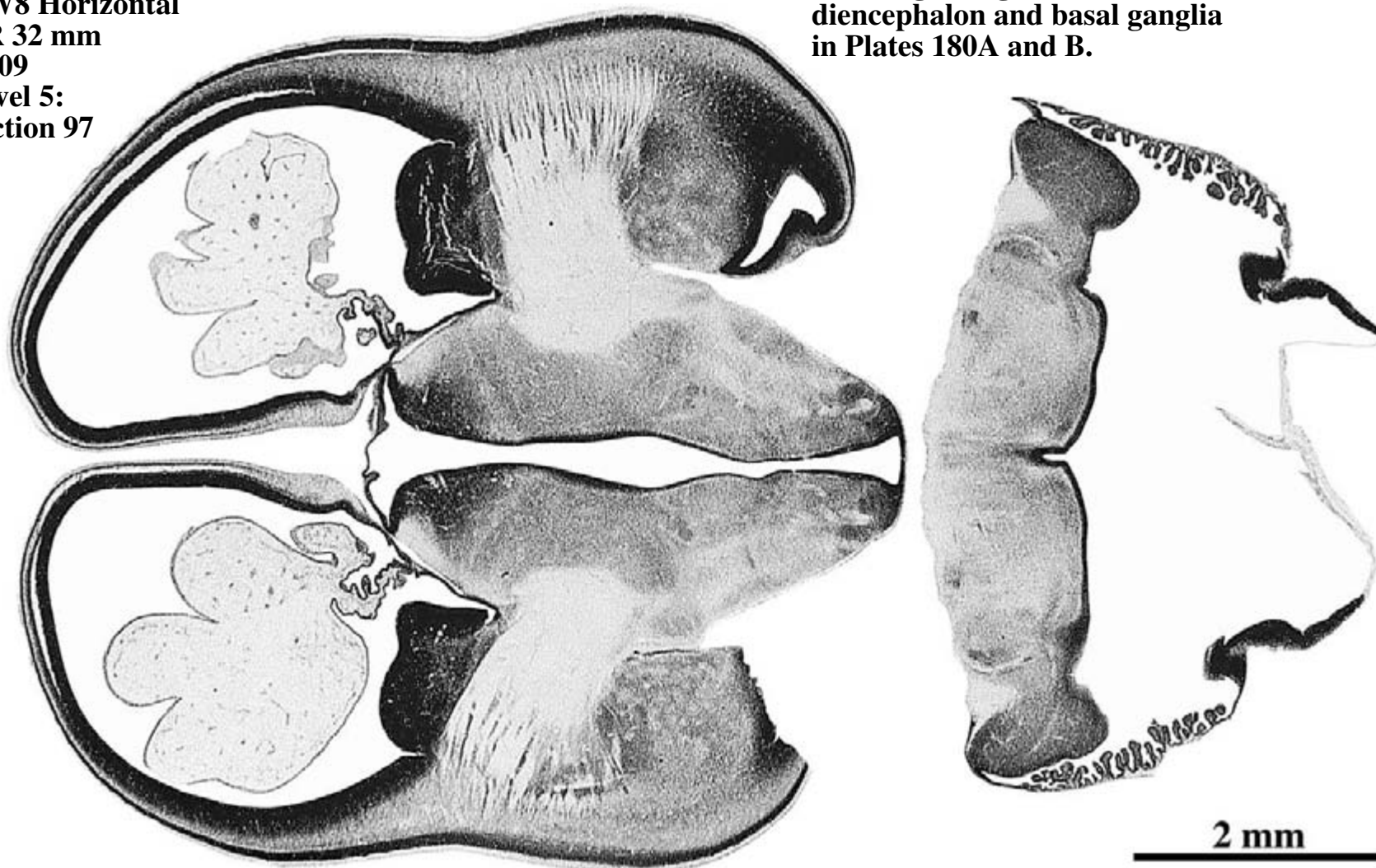
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 174A

GW8 Horizontal
CR 32 mm
C609
Level 5:
Section 97

See a high-magnification view of the
diencephalon and basal ganglia
in Plates 180A and B.

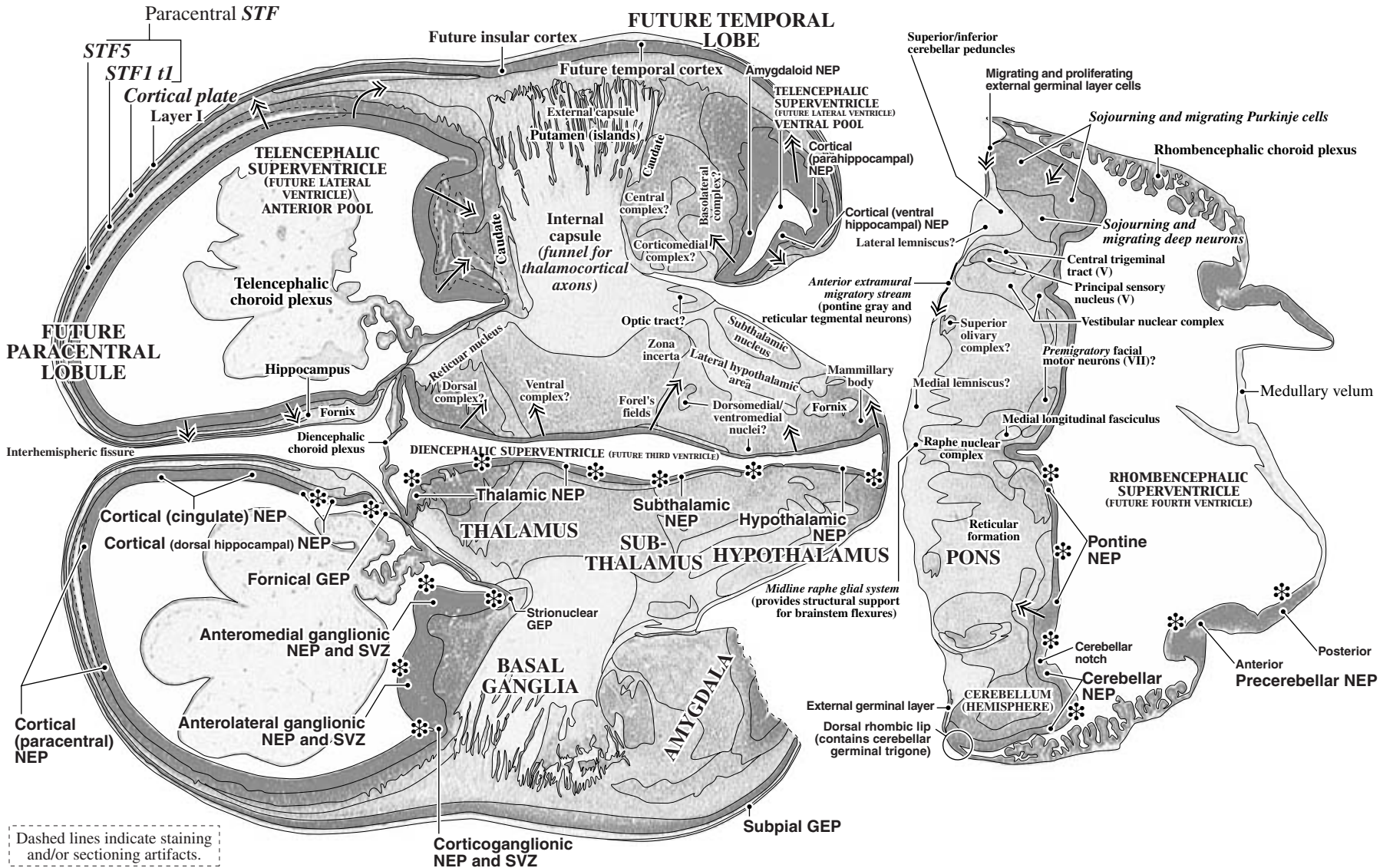


LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

STF1 Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.

STF5 Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 174B



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

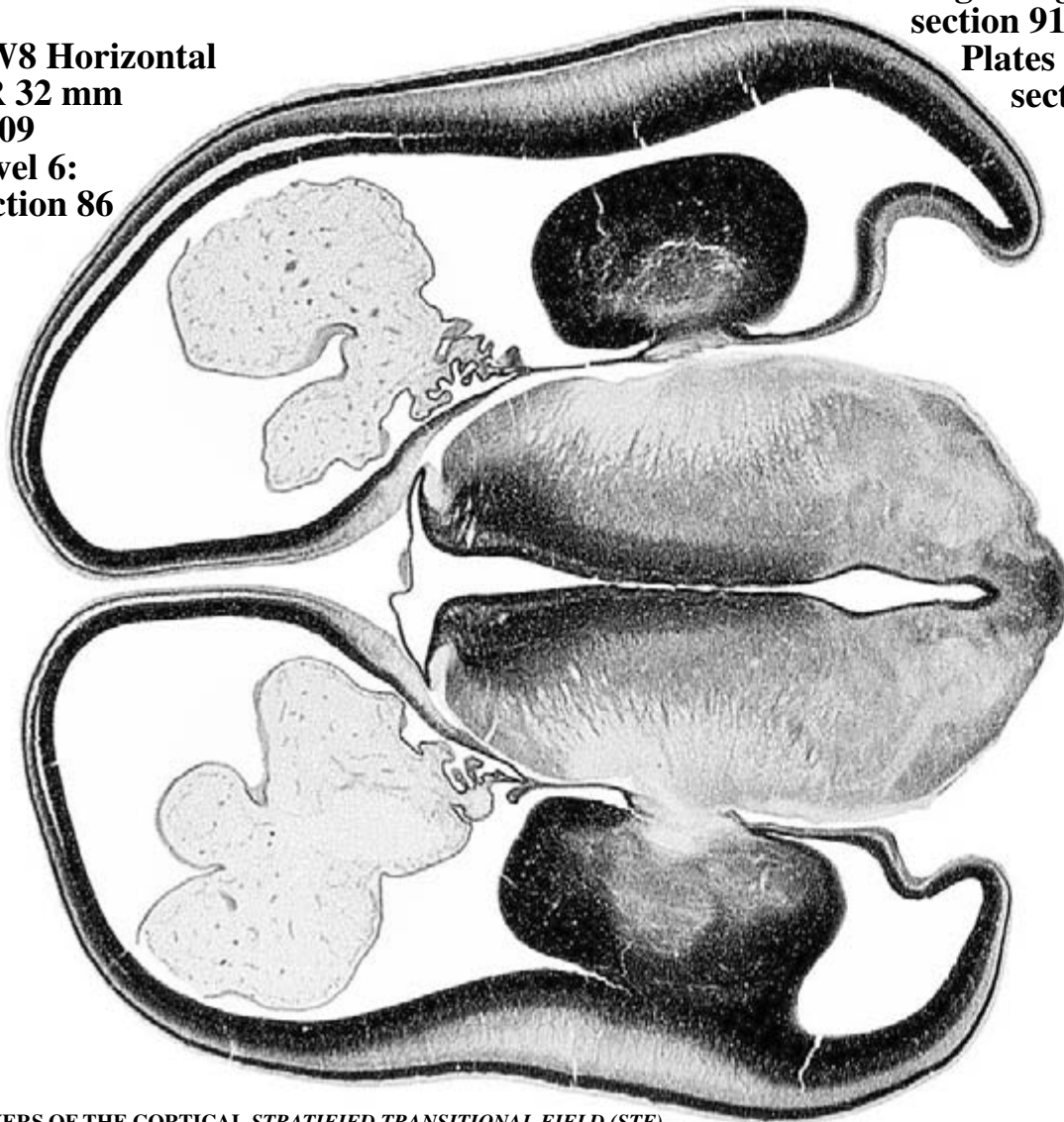
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 175A

**GW8 Horizontal
CR 32 mm
C609
Level 6:
Section 86**

See high-magnification views of the diencephalon from section 91 in Plates 181A and B, from this section in Plates 182A and B, and of the hypothalamus from section 92 in Plates 185A and B.

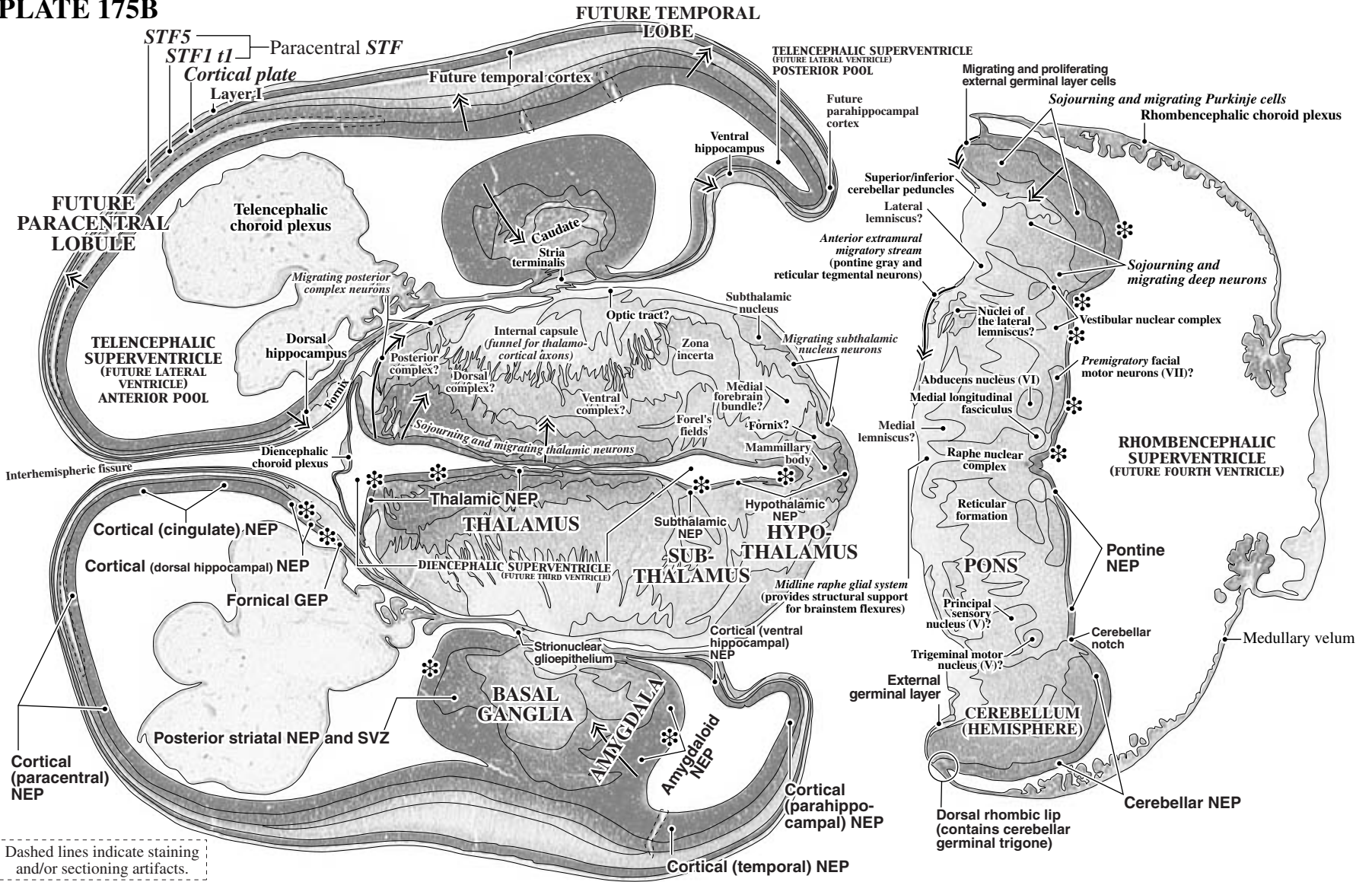


2 mm

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 175B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

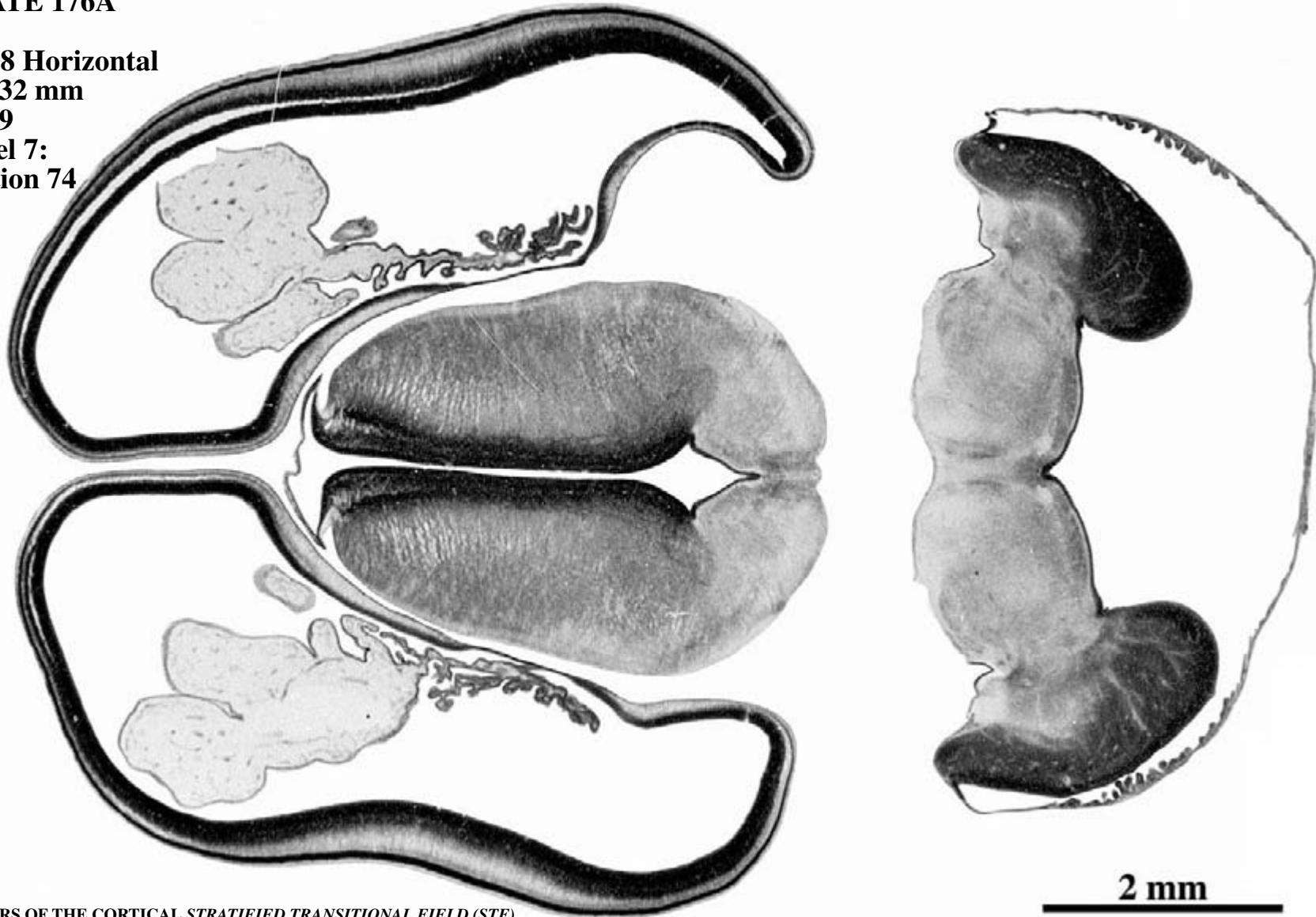
ABBREVIATIONS:
GEP - Glioeptithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 176A

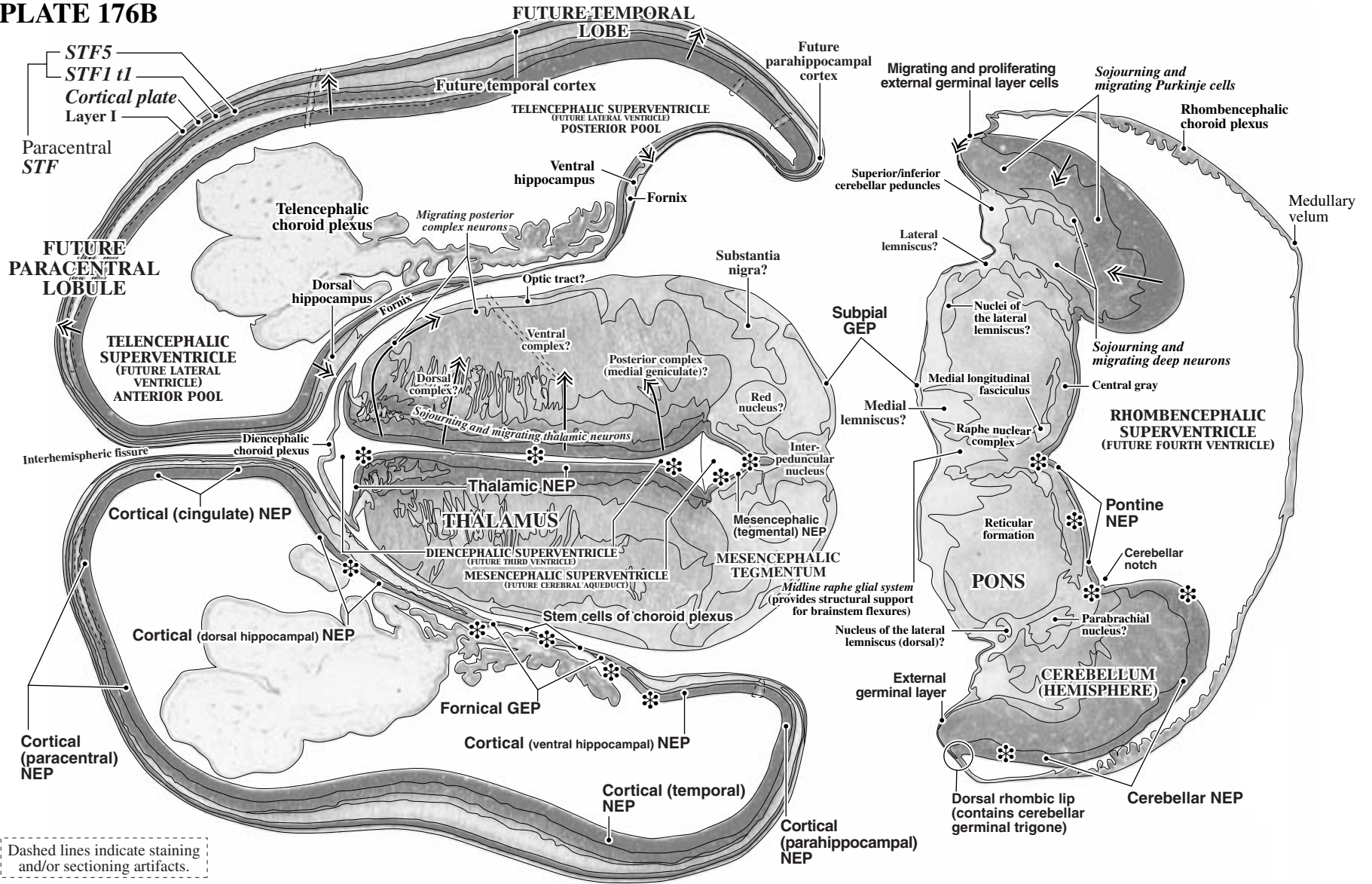
**GW8 Horizontal
CR 32 mm
C609
Level 7:
Section 74**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 176B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

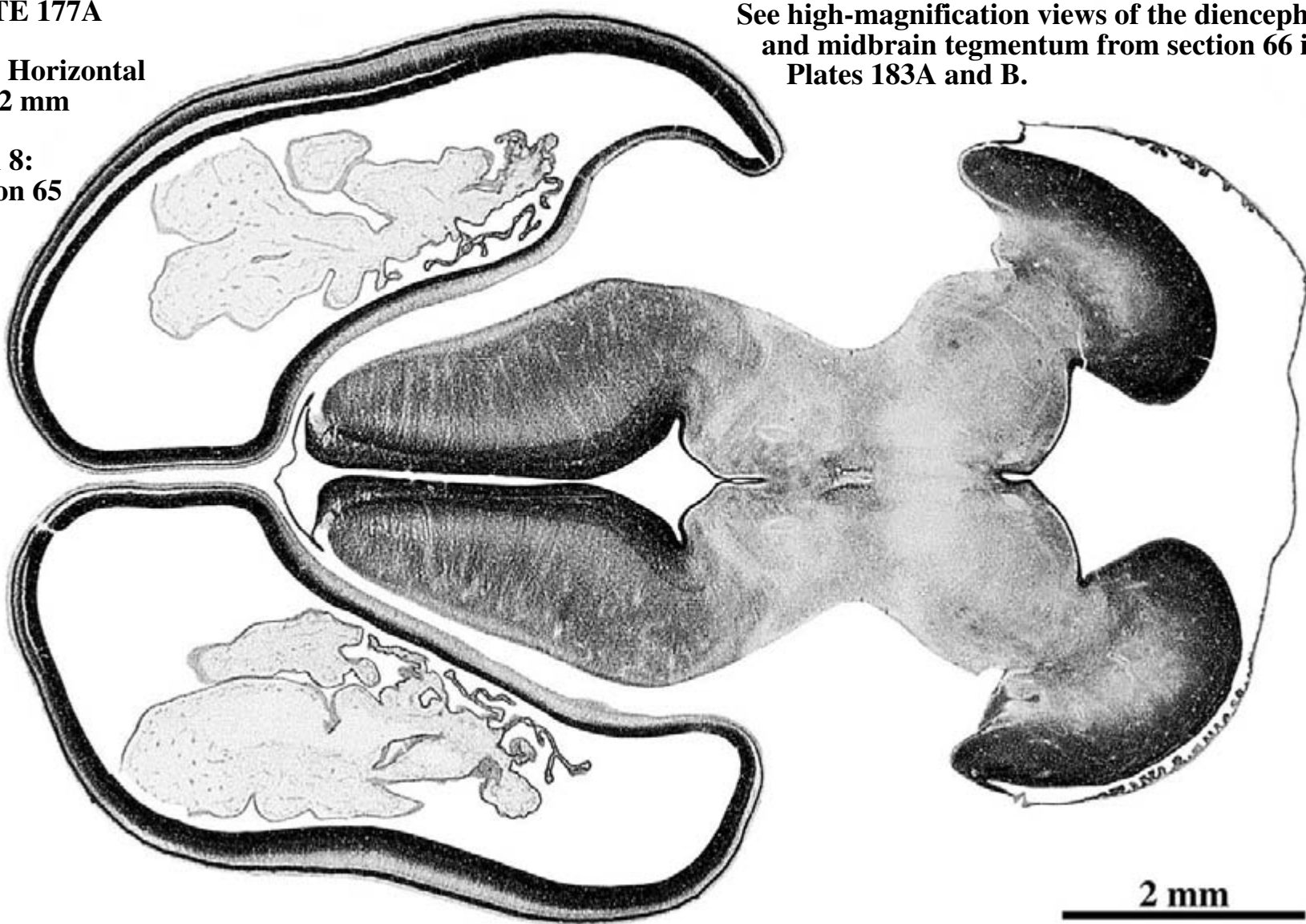
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 177A

**GW8 Horizontal
CR 32 mm
C609
Level 8:
Section 65**

See high-magnification views of the diencephalon and midbrain tegmentum from section 66 in Plates 183A and B.

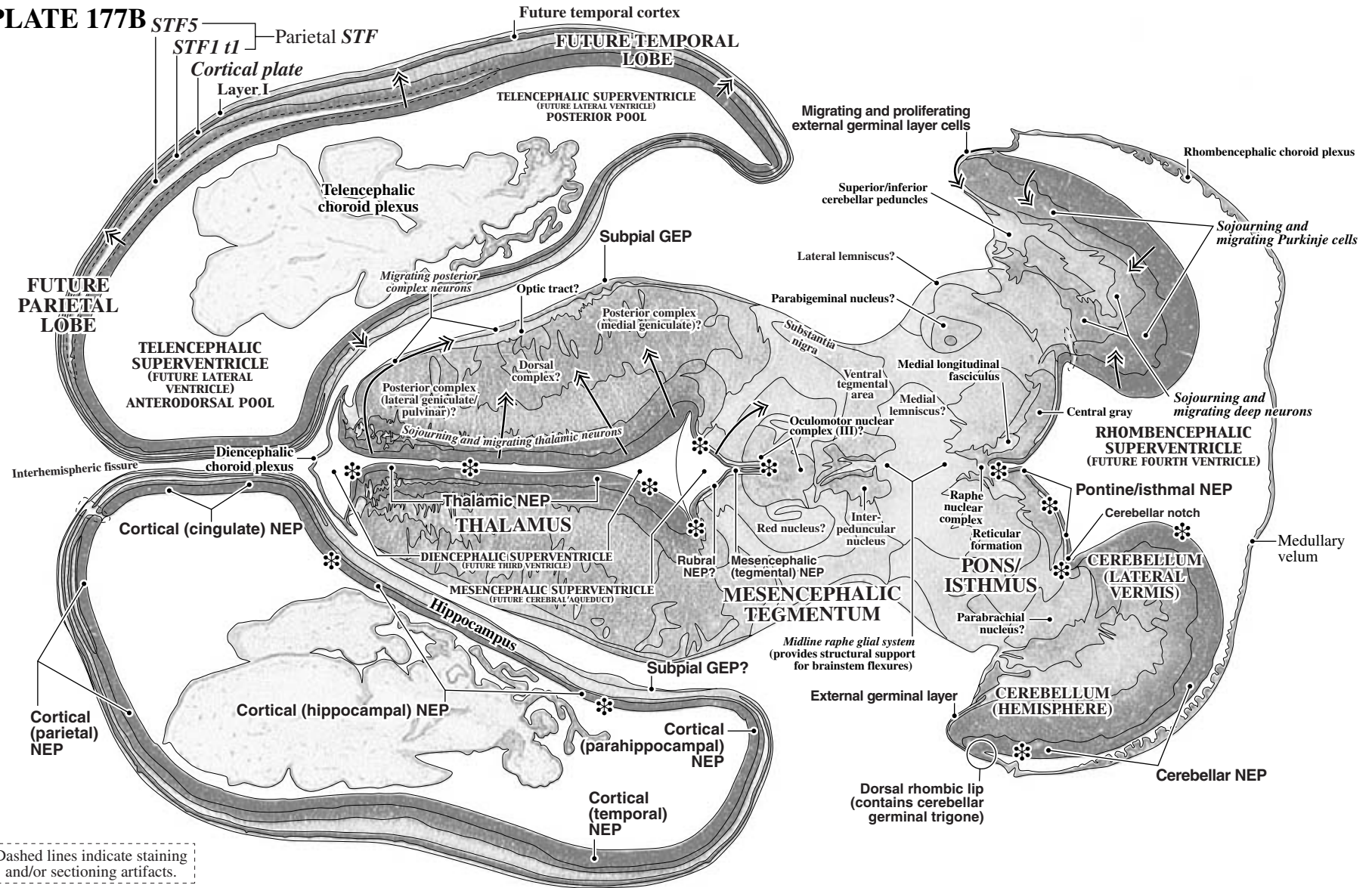


2 mm

LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 177B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

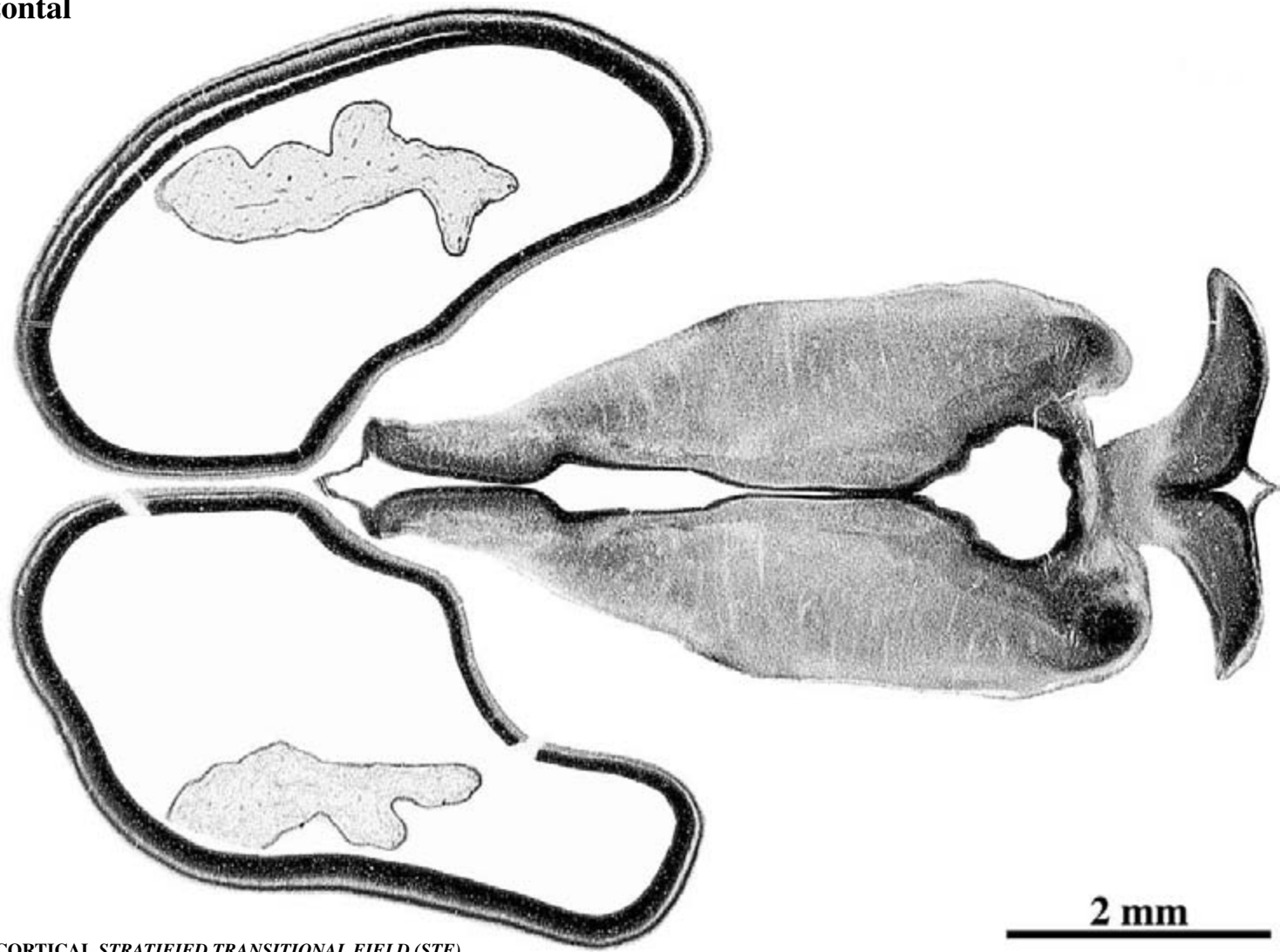
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

PLATE 178A

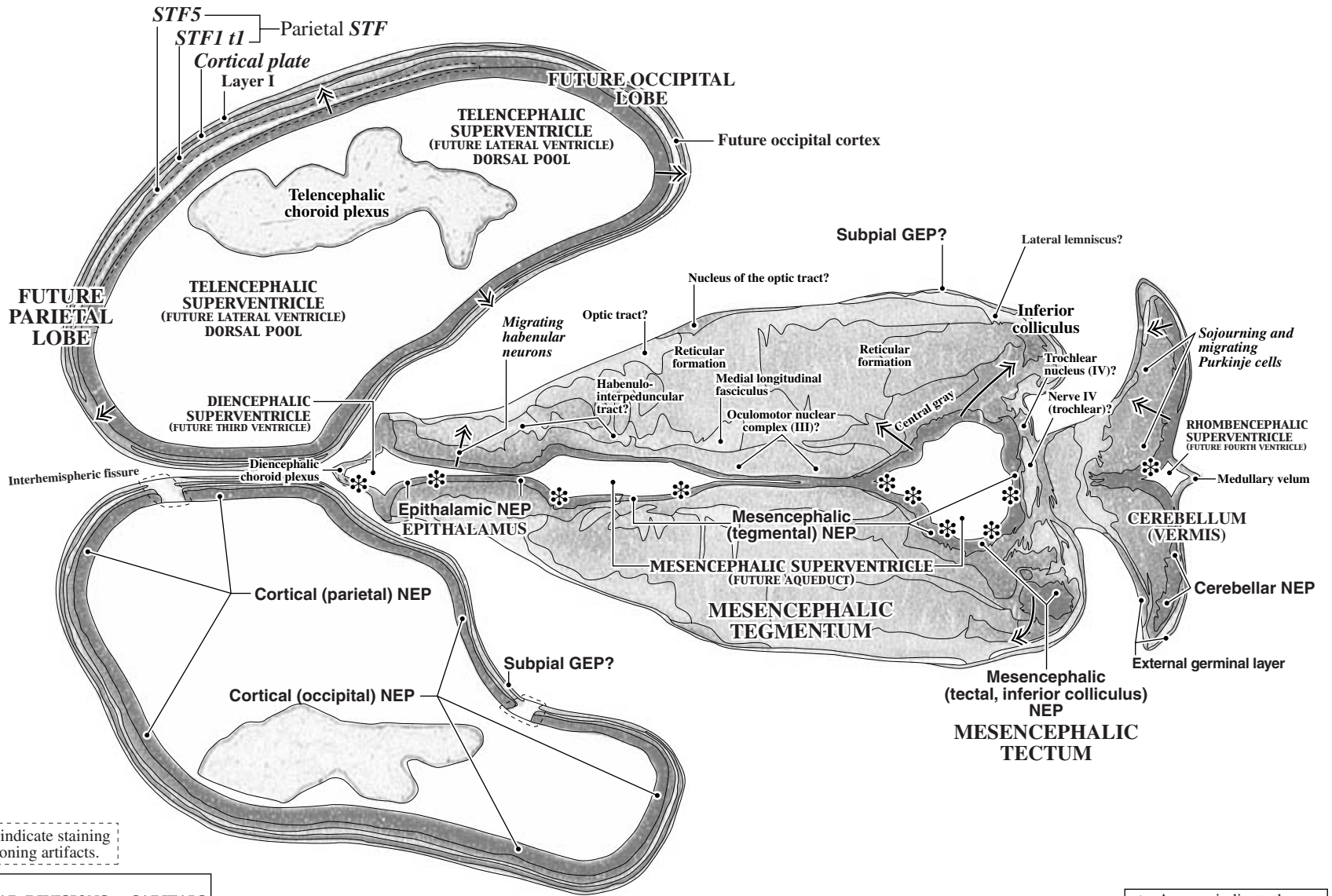
**GW8 Horizontal
CR 32 mm
C609
Level 9:
Section 45**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 178B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

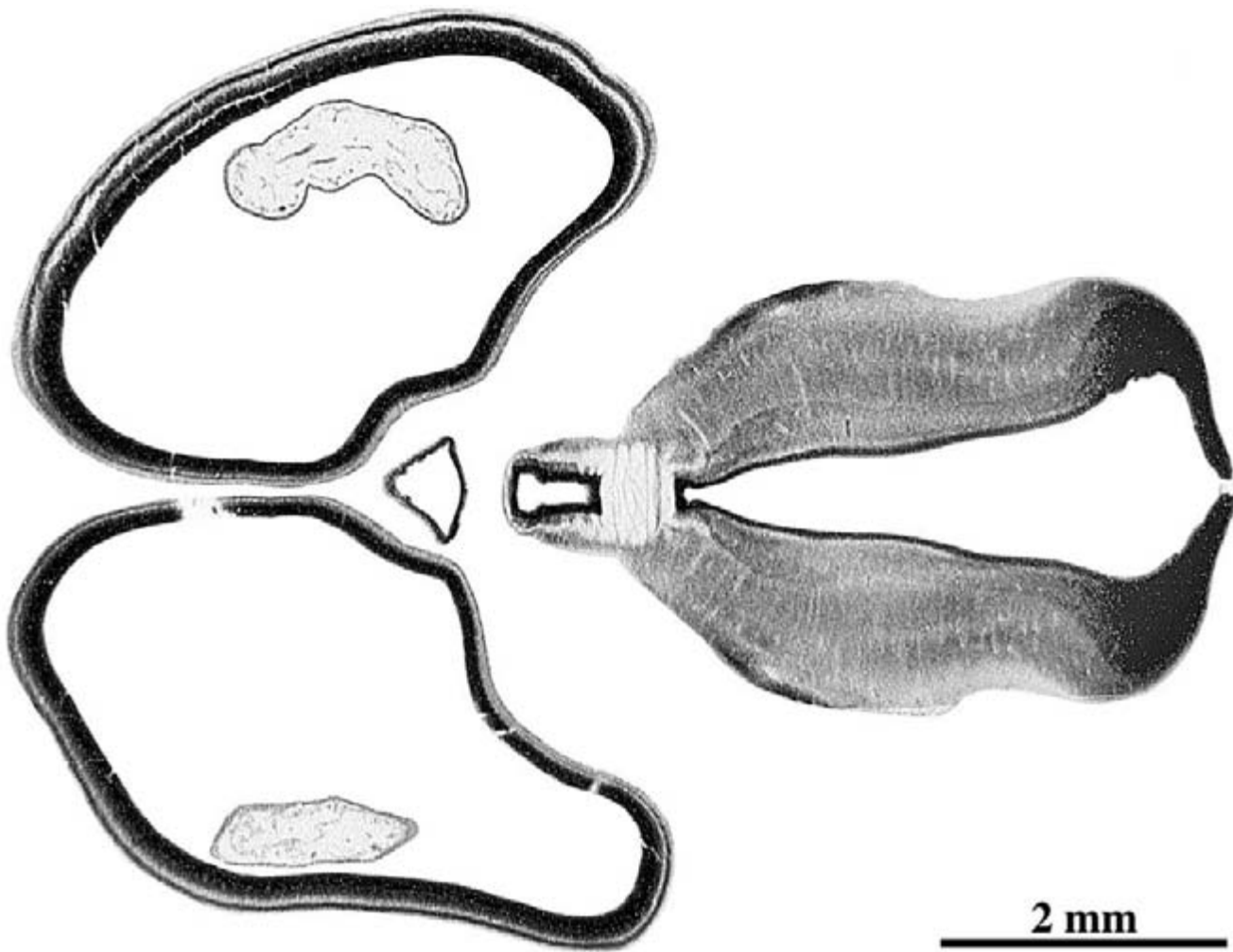
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 179A

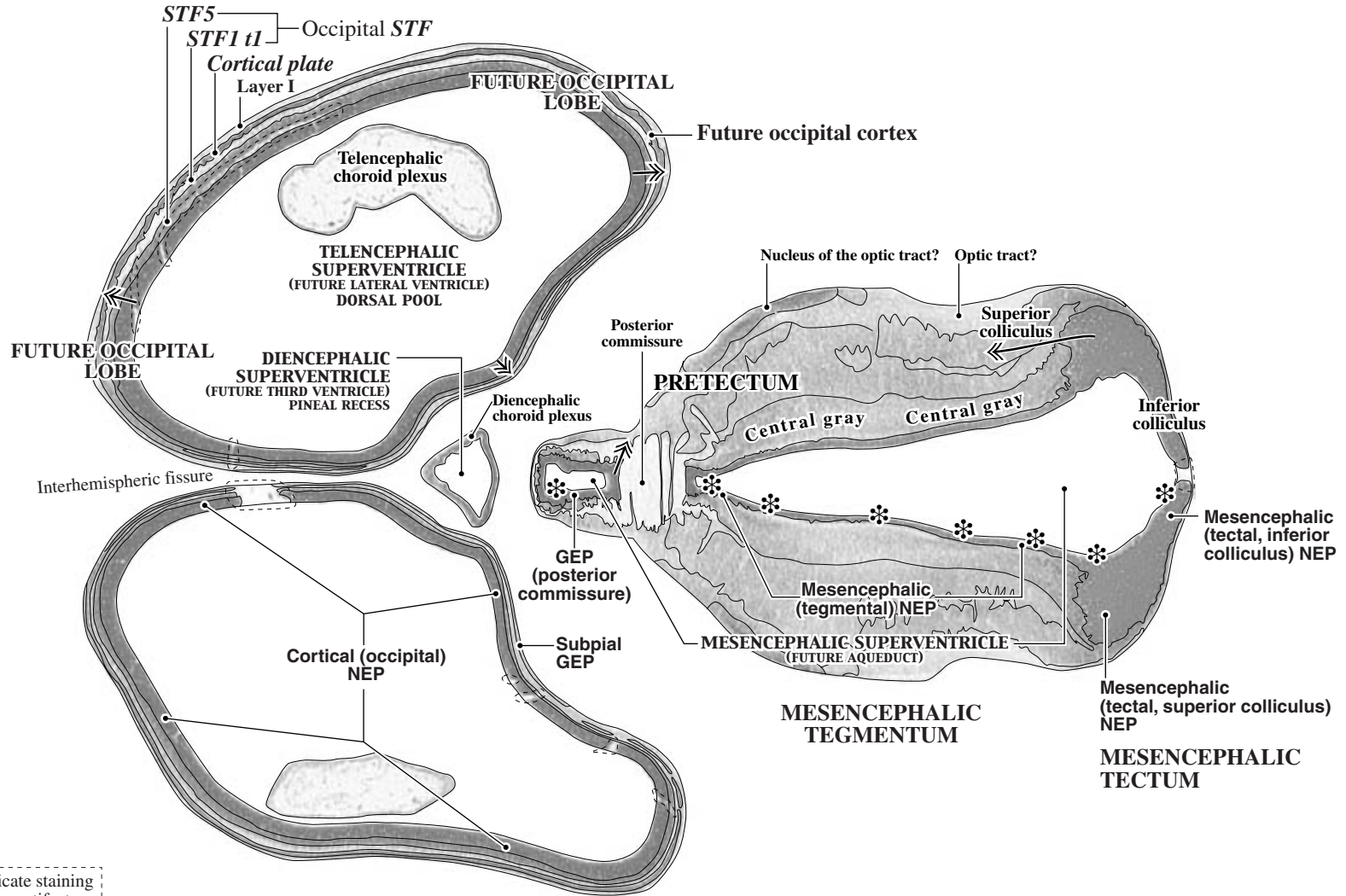
**GW8 Horizontal
CR 32 mm
C609
Level 10:
Section 36**



LAYERS OF THE CORTICAL STRATIFIED TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*11*) when many cells are migrating through it, followed by a late stage (*12*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

PLATE 179B



Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Gliopithelium
NEP - Neuroepithelium

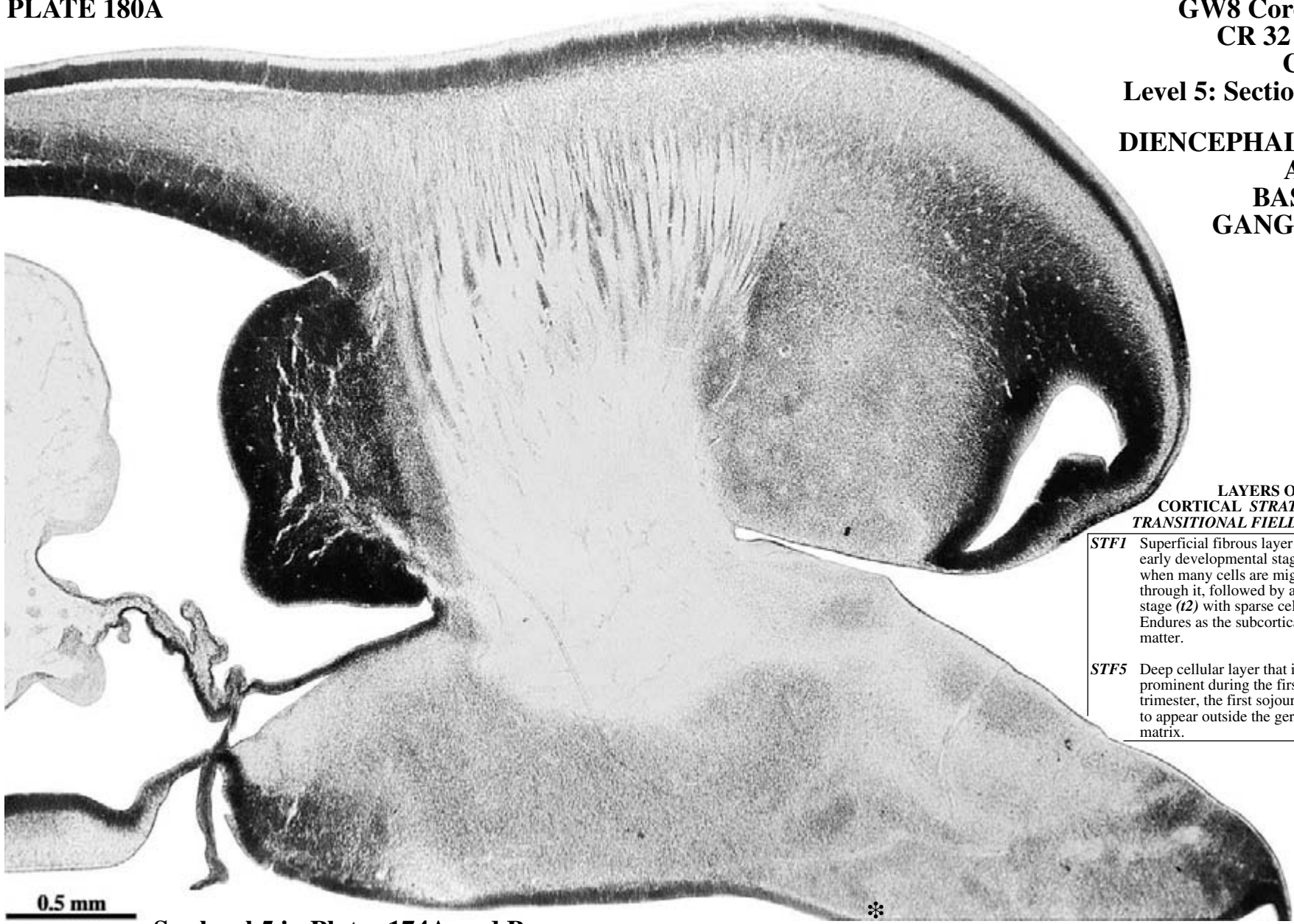
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 180A

GW8 Coronal
CR 32 mm
C609
Level 5: Section 97

DIENCEPHALON
AND
BASAL
GANGLIA



LAYERS OF THE
CORTICAL STRATIFIED
TRANSITIONAL FIELD (STF)

- STF1** Superficial fibrous layer with an early developmental stage (*t1*) when many cells are migrating through it, followed by a late stage (*t2*) with sparse cells. Endures as the subcortical white matter.
- STF5** Deep cellular layer that is prominent during the first trimester, the first sojourn zone to appear outside the germinal matrix.

0.5 mm

See level 5 in Plates 174A and B.

*

PLATE 180B

Dashed lines indicate staining and/or sectioning artifacts.

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

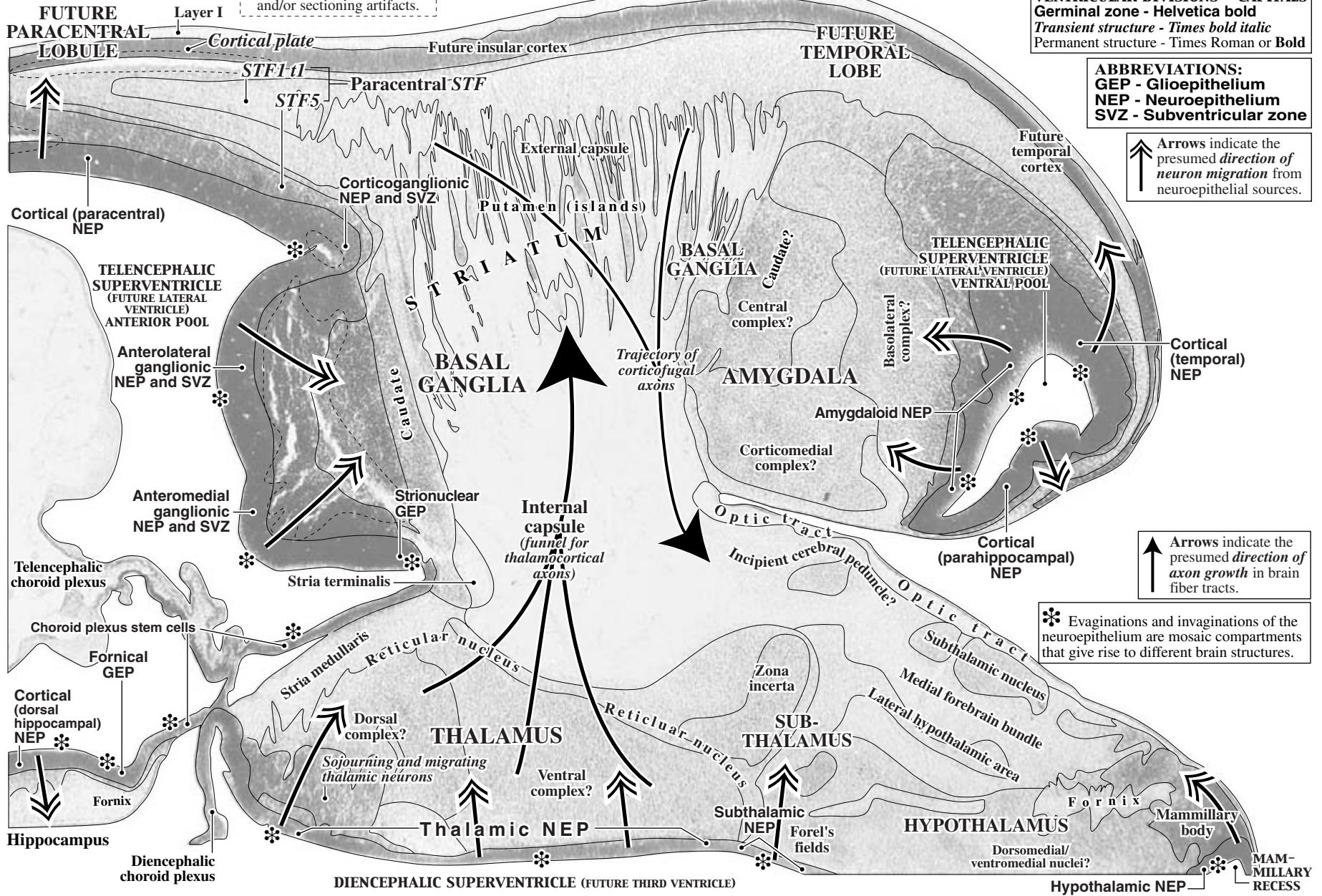
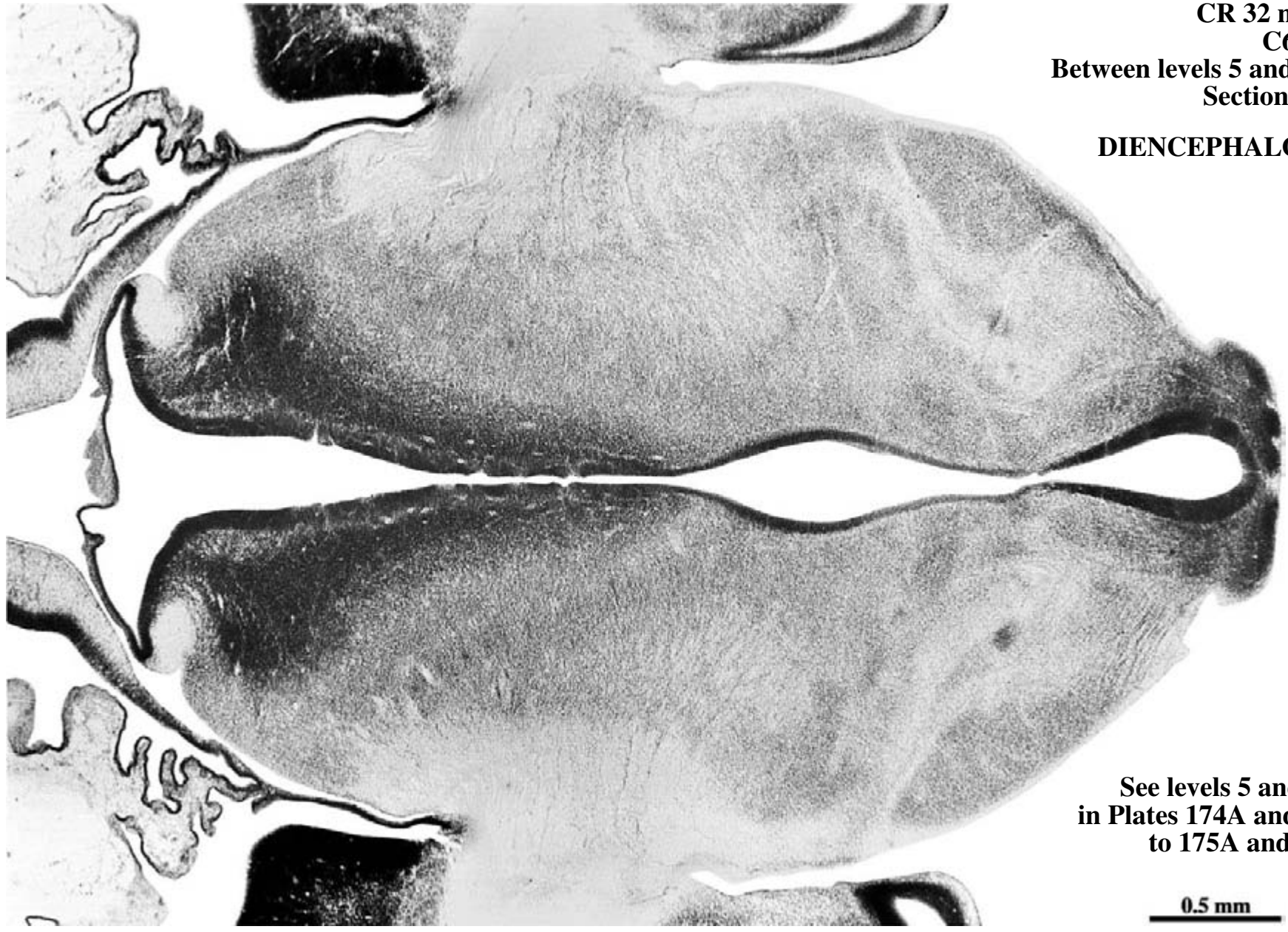


PLATE 181A

**GW8 Coronal
CR 32 mm
C609
Between levels 5 and 6:
Section 91**

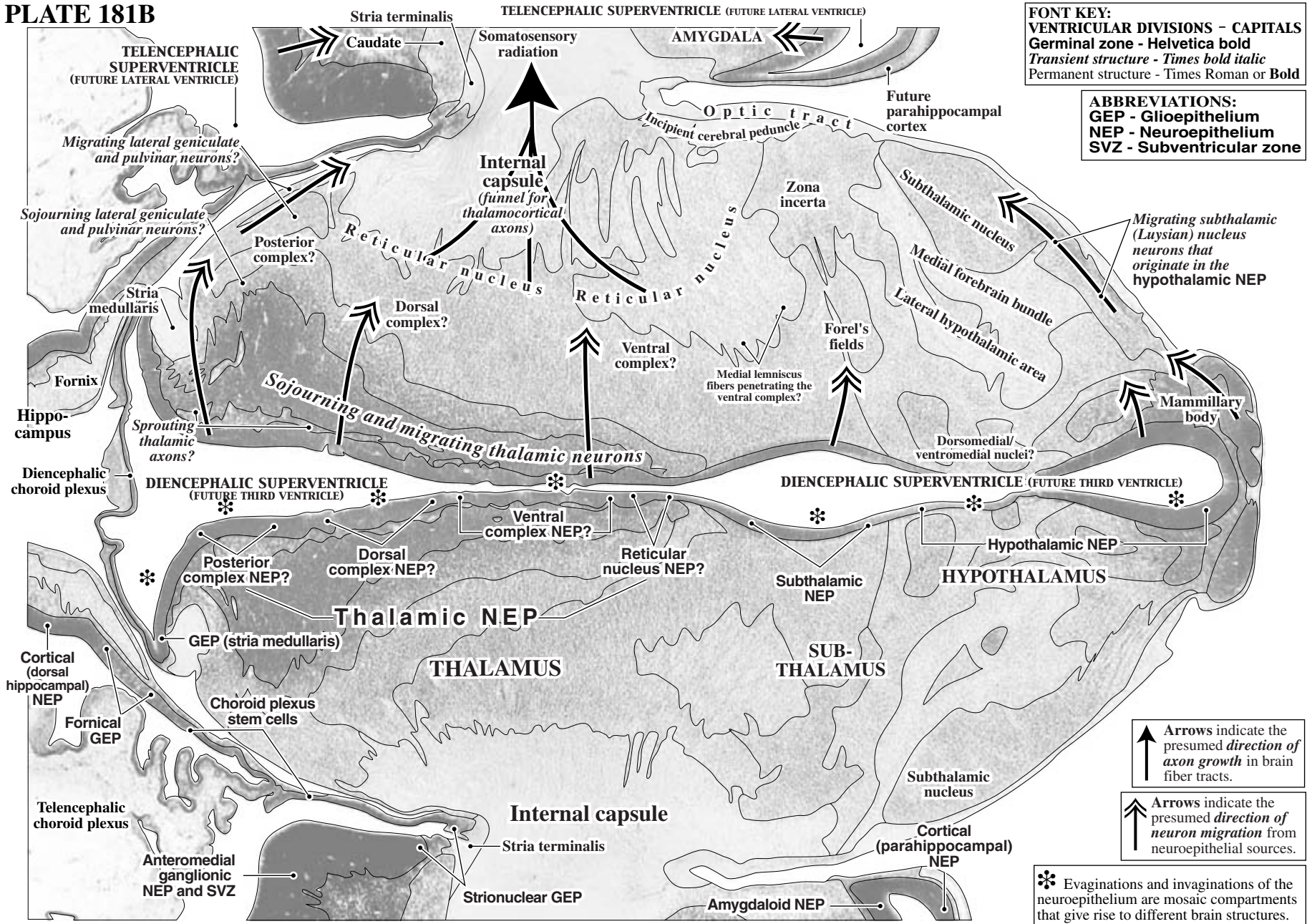
DIENCEPHALON



**See levels 5 and 6
in Plates 174A and B
to 175A and B.**

0.5 mm

PLATE 181B



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

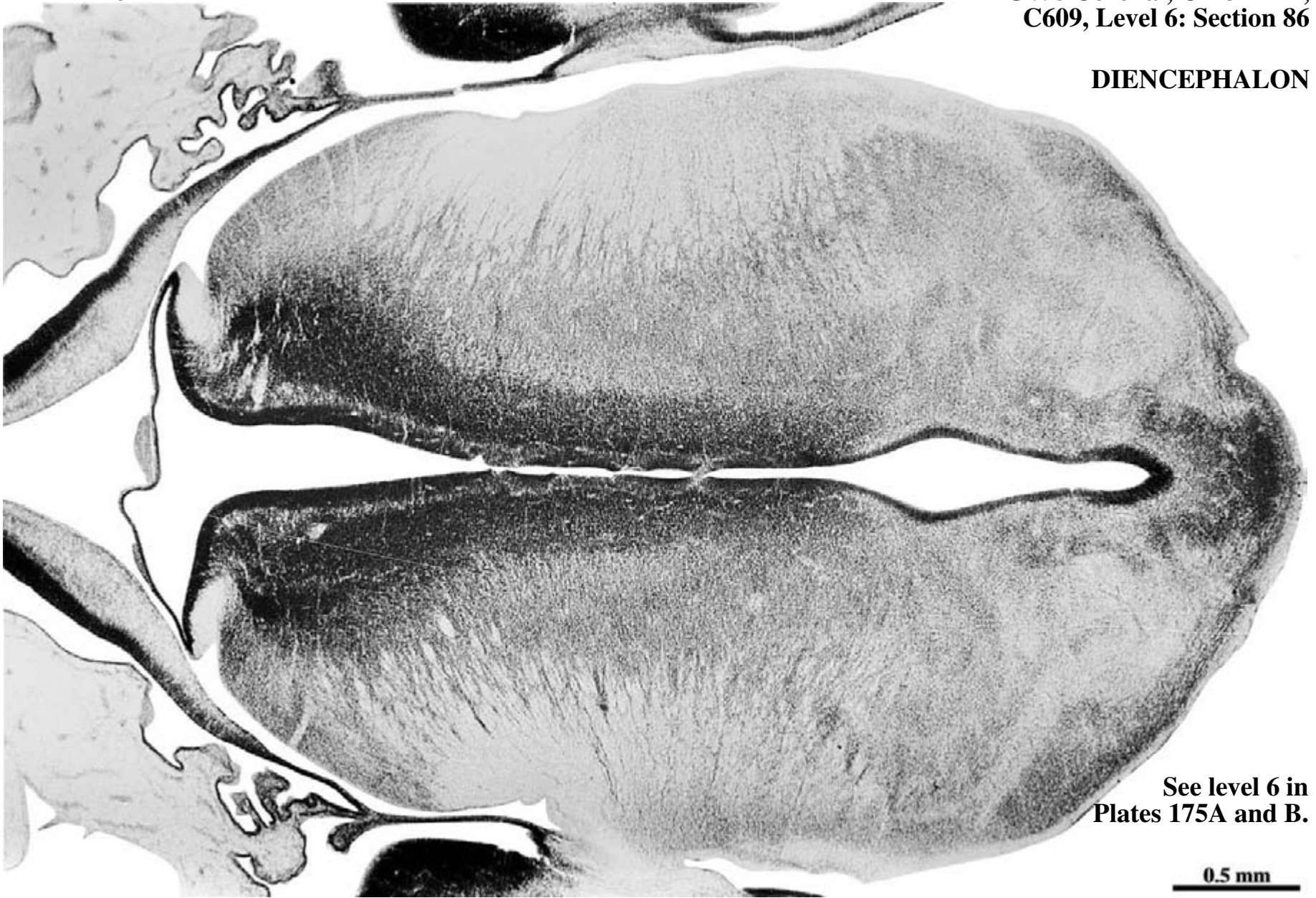
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 182A

**GW8 Coronal, CR 32 mm,
C609, Level 6: Section 86**

DIENCEPHALON



**See level 6 in
Plates 175A and B.**

0.5 mm

PLATE 182B

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

TELENCEPHALIC SUPERVENTRICLE
 (FUTURE LATERAL VENTRICLE)

Caudate Stria terminalis

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

↑ Arrows indicate the presumed *direction of axon growth* in brain fiber tracts.

↑ Migrating subthalamic (Luysian) nucleus neurons that originate in the hypothalamic NEP

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

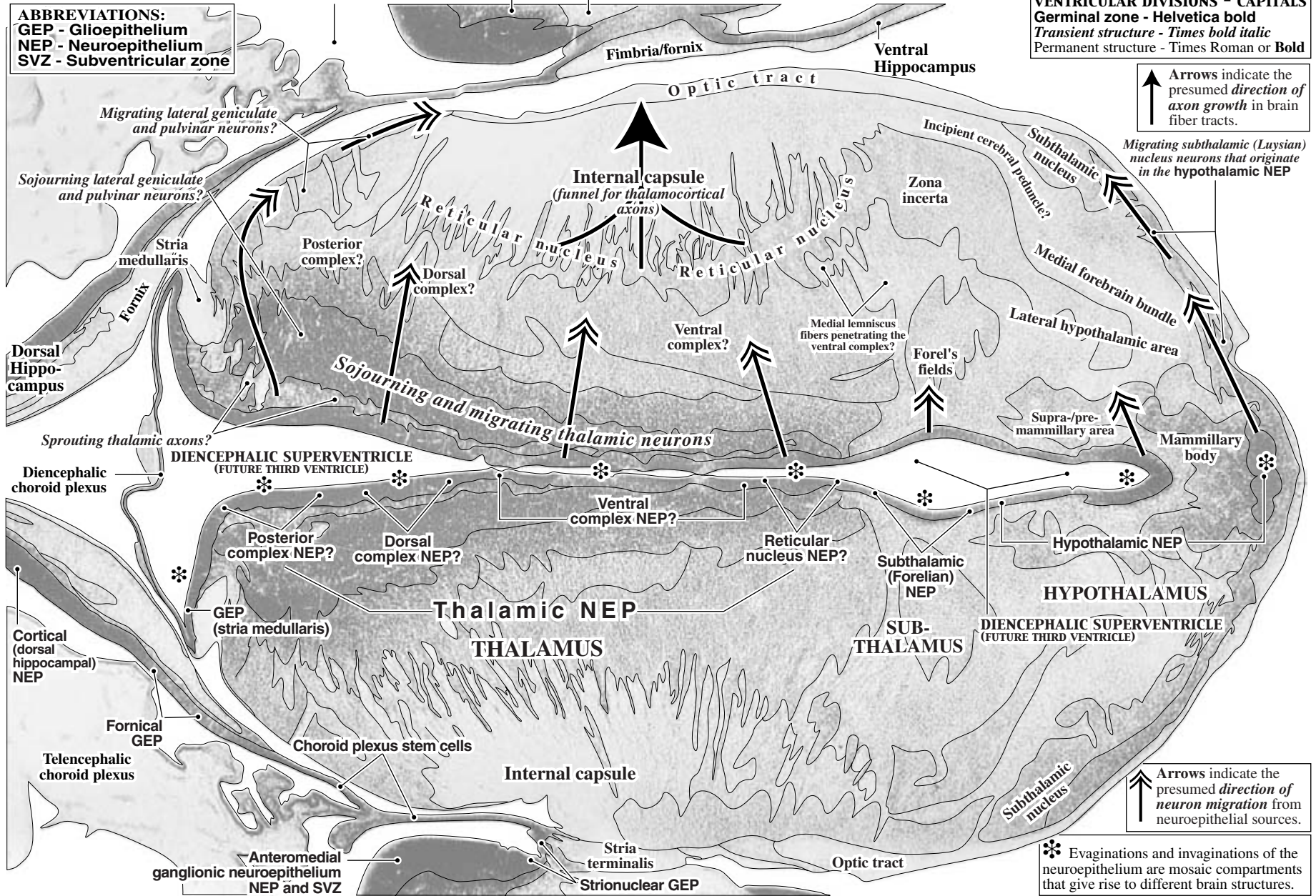
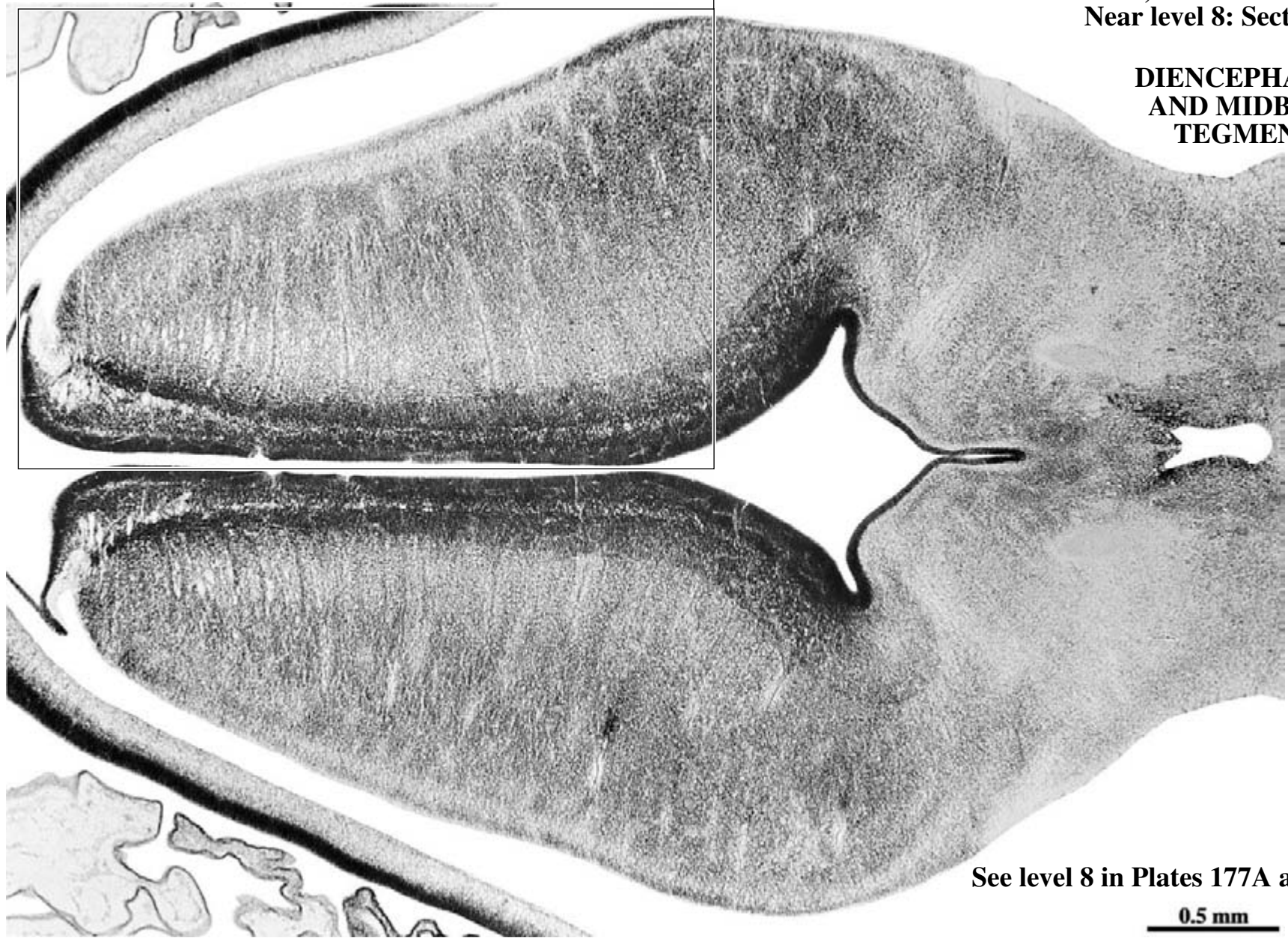


PLATE 183A

Enlarged in Plates 184A and B.

**GW8 Coronal, CR 32 mm, C609
Near level 8: Section 66**

**DIENCEPHALON
AND MIDBRAIN
TEGMENTUM**



See level 8 in Plates 177A and B.

0.5 mm

PLATE 183B

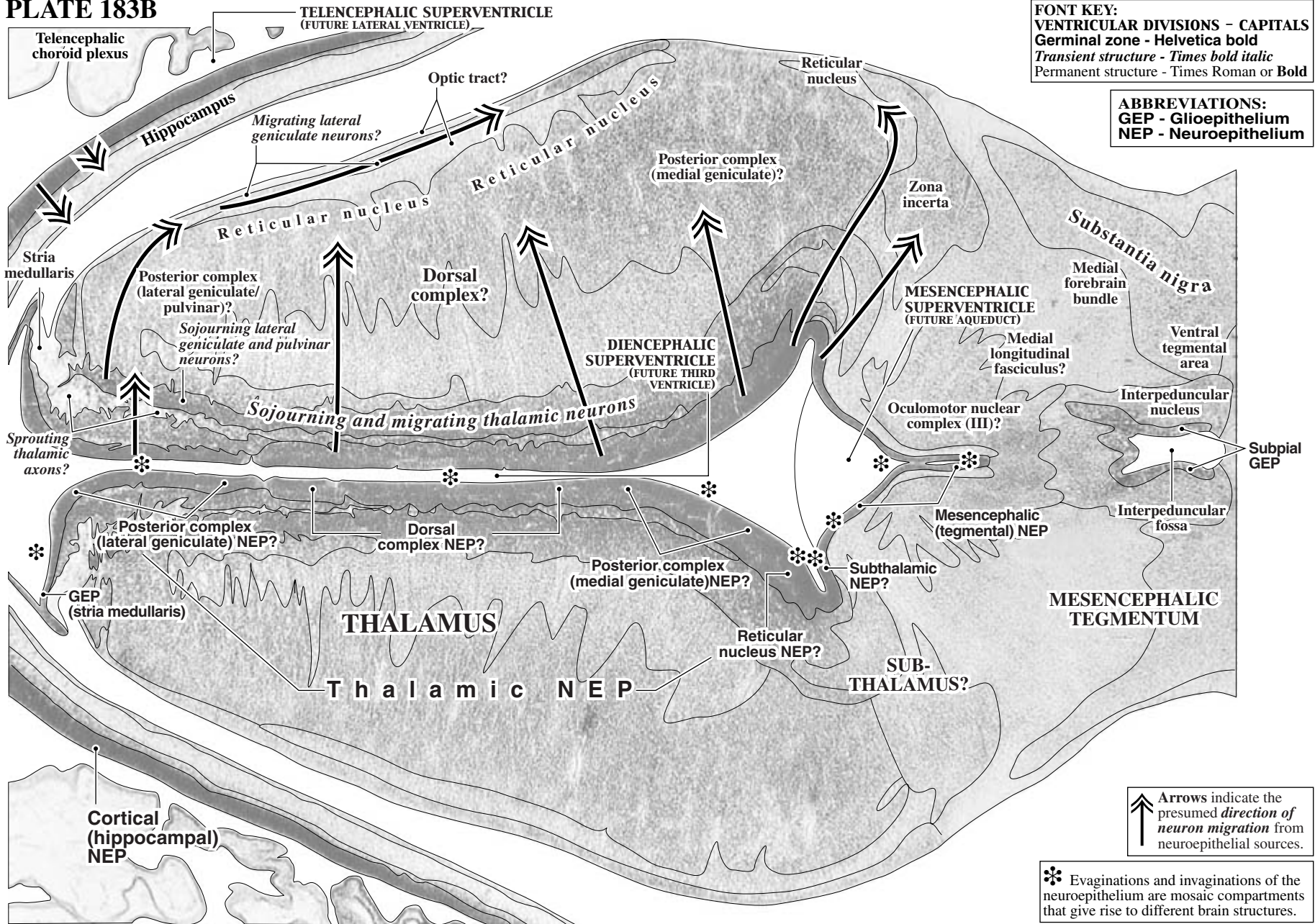
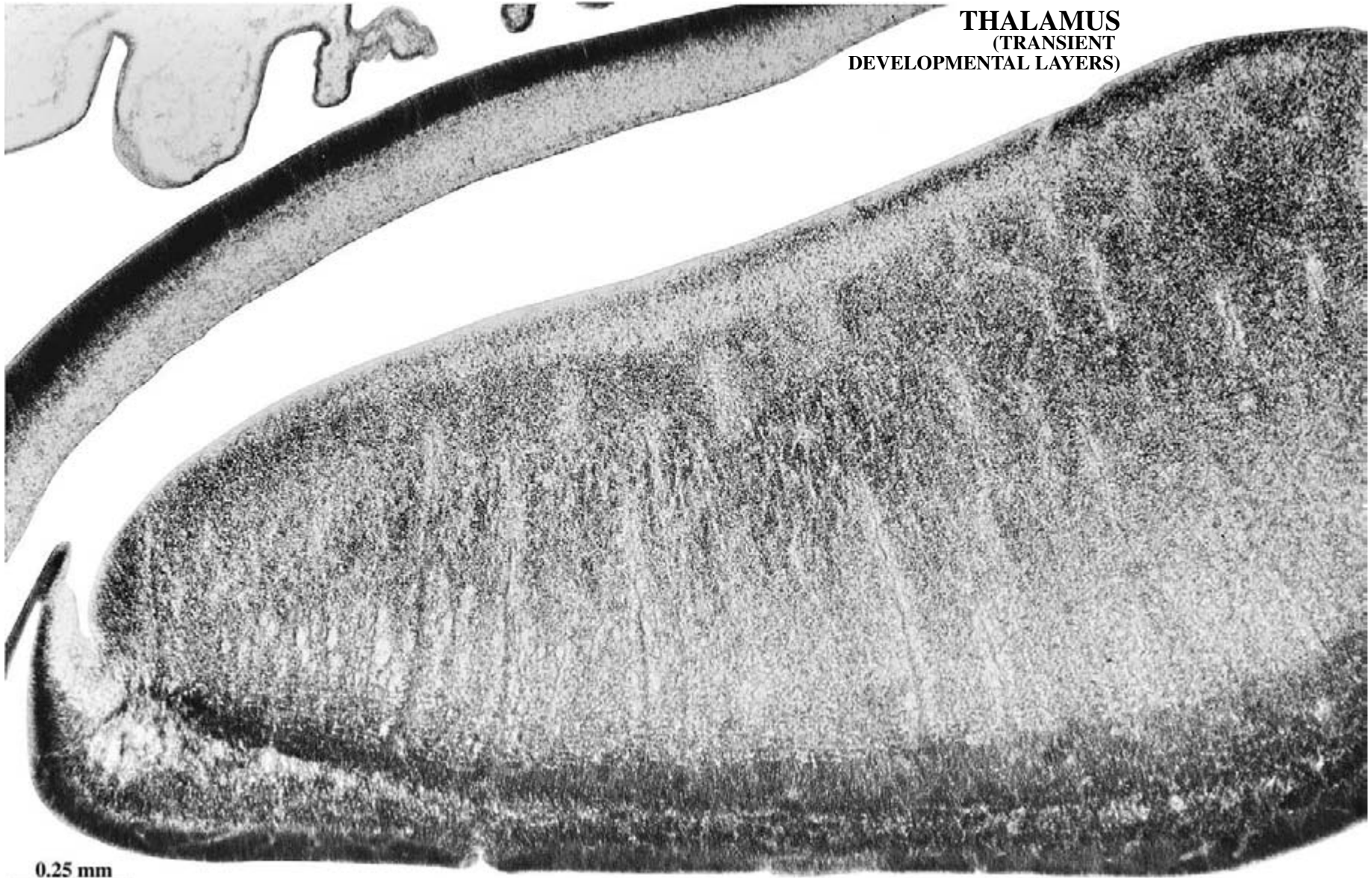


PLATE 184A

GW8 Coronal, CR 32 mm, C609, Near level 8: Section 66



**THALAMUS
(TRANSIENT
DEVELOPMENTAL LAYERS)**

0.25 mm

See level 8 in Plates 177A and B.

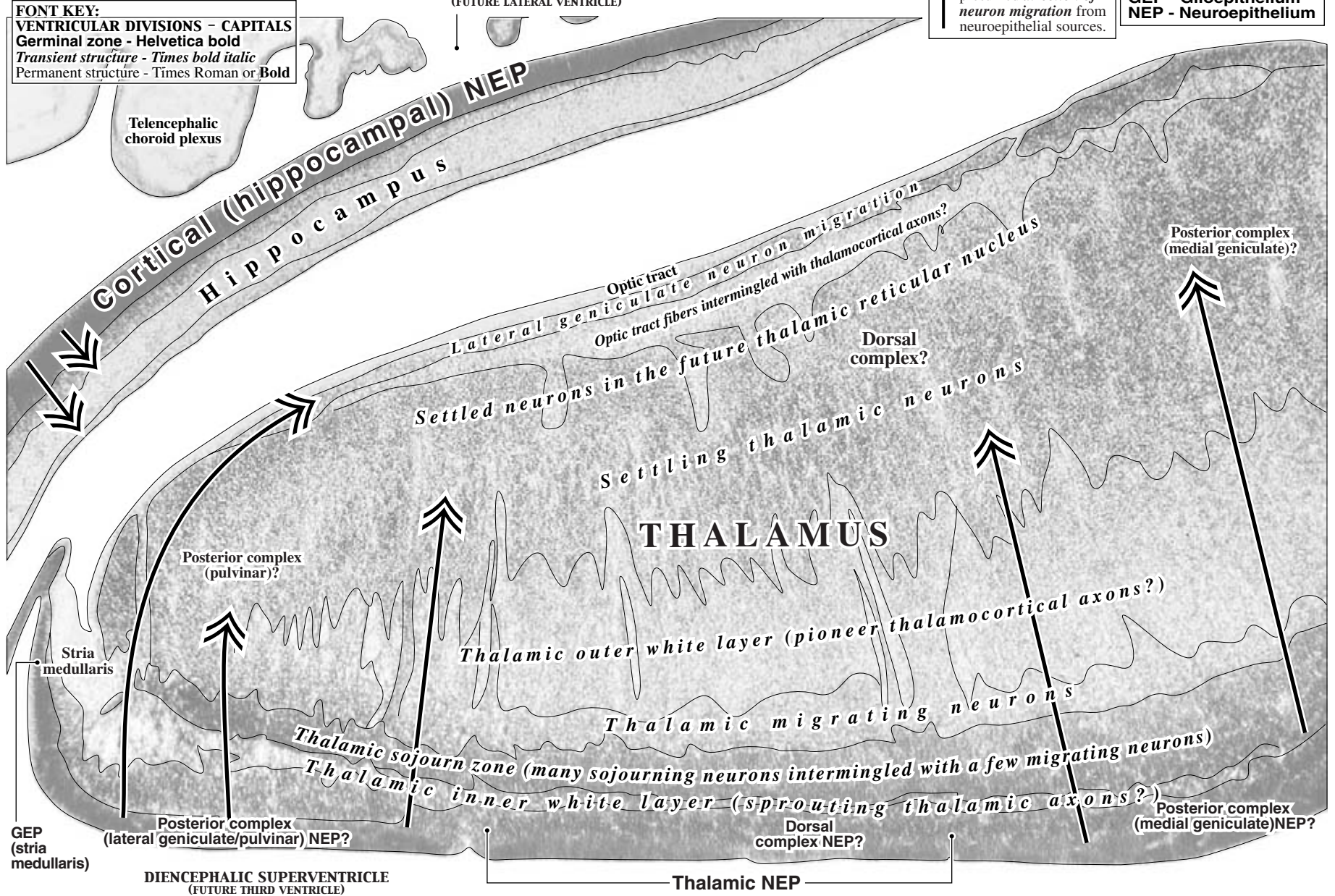
PLATE 184B

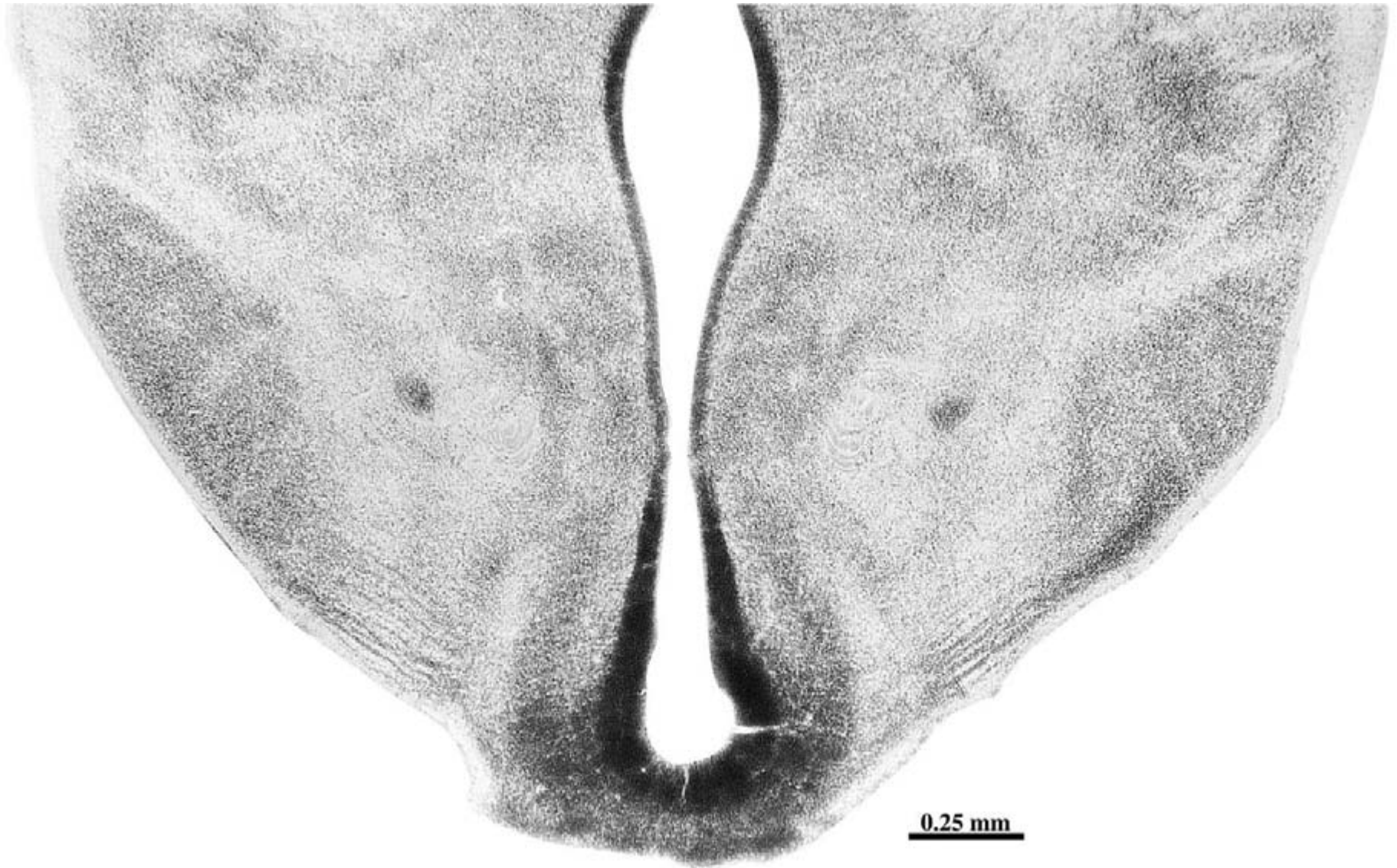
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

TELENCEPHALIC SUPERVENTRICLE
 (FUTURE LATERAL VENTRICLE)

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

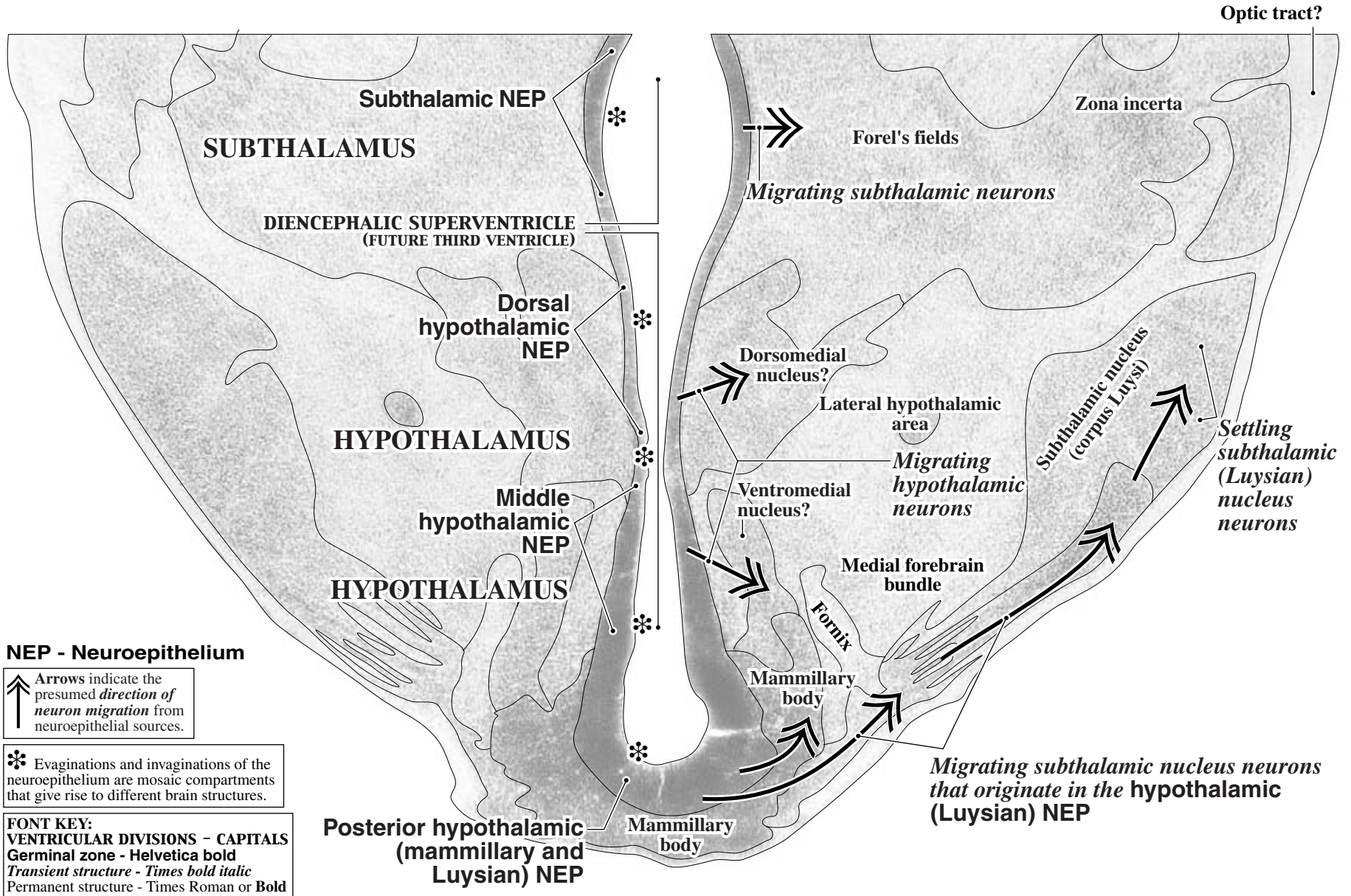
ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium





See levels 5 and 6 in Plates 174A and B to 175A and B.

PLATE 185B



PART X: GW7.5 CORONAL

This is specimen number 966 in the Carnegie Collection, designated here as C966. A normal female fetus with a crown-rump length (CR) of 23 mm was collected in 1914. The fetus is estimated to be at gestational week (GW) 7.5. The entire fetus was fixed in bichloric acetic acid, embedded in celloidin, cut transversely in 40 μm sections, and was stained with aluminum cochineal. The histological preservation of this specimen is excellent, and the sections are cut nearly perfectly bilaterally. Several years ago, an excellent 3-D reconstruction of the brain and upper cervical spinal cord was done by piecing together cardboard cutouts of the brain outlines in each section and then gluing them together; the rhombencephalic superventricle was not included in that reconstruction. A photograph of that model (which is still part of the Carnegie Collection today) shows us the exact location and cutting plane of C966's sections (**Figure 9**). Like most of the specimens in this Volume, the sections are not cut exactly in one plane, but C966's sections are much closer to the coronal than the horizontal plane. Photographs of 21 sections (**Levels 1-21**) of the brain in the head are shown in **Plates 186-206**. Our computer-aided three-dimensional reconstructions of the brain, ventricles (including the rhombencephalic superventricle), and selected neuroepithelial components are shown on the cover and in **Figures 10-19**.

C966 is considerably less mature than the GW8 specimens. The *superventricles* are large in the centers of all brain structures, especially in the telencephalon and rhombencephalon. Even though the diencephalic superventricle is approaching a slit-like shape, it is wider than that in the GW8 specimens, and the mesencephalic superventricle forms a more balloon-like expansion beneath the rudimentary tectum. Like the GW8 specimens, the respective thicknesses of the neuroepithelium (NEP) and parenchyma are keys to determining the degree of maturation of various brain structures.

The parenchyma is thick and bordered by a thin NEP in the medulla and pons, indicating that many neurons have been generated here, but the production of late-generated neurons continues. There are layers of dense cells adjacent to the lateral pontine NEP where vestibular nuclear neurons and trigeminal nuclear neurons may be sojourning prior to migration and settling. There is a larger accumula-

tion of presumptive facial motor neurons near the midline NEP in the pontomedullary trench and some are migrating toward the indistinct facial motor nucleus. The *precerebellar neuroepithelium* in the medulla is thicker and generating more precerebellar neurons than at GW8; many neurons are entering the inferior olive after migrating in the *posterior intramural migratory stream*, but the anterior and posterior extramural migratory streams are absent; that confirms neurogenetic data in rats that the inferior olive contains the earliest-generated precerebellar neurons. The cerebellar NEP is thick and difficult to distinguish from an adjacent dense sojourn zone in the cerebellar parenchyma, called *cerebellar transitional field (CTF) 6*. The remaining *CTF* has alternating layers of cells and fibers (*CTF1-5*). The *external germinal layer (egl)* is completely absent. If one can extrapolate from data on cerebellar neurogenesis in rats, the human cerebellar NEP by GW7.5 has generated all of the deep neurons and is now producing "middle-aged" Purkinje cells; the oldest Purkinje cells are sojourning in *CTF6*. Both the mesencephalic tegmental NEP and the isthmal NEP are nearly the same thickness as the pontine and medullary NEPs, but dense sojourn zones of young neurons are more obvious in the adjacent parenchyma. The superficial border of a thick mesencephalic tectal NEP is difficult to distinguish from dense wavefronts of young neurons extending into a thin parenchyma. The majority of neurons in both the superior and inferior colliculi have not yet been generated by GW7.5.

The diencephalic NEP is thicker at GW7.5 than at GW8, and the thin parenchyma is filled with dense zones of sojourning and migrating neurons. There is no internal capsule, but some pioneer axons are accumulating in inner and outer white layers in the thalamic parenchyma. The telencephalic NEP is thick in all areas, and the oldest basal telencephalic and basal ganglionic neurons are settling in the thick parenchyma. Most neurons in the septum and striatum have yet to be generated. In the cerebral cortex, the NEP is bordered by a thin primordial plexiform layer that contains the oldest cortical neurons (Cajal-Retzius cells) and subplate neurons; there is no cortical plate. The cortical NEP is expanding and increasing its number of neural stem cells as the telencephalic superventricle grows; nearly all cortical neurons in layers II-VI have still to be generated.

GW7.5 CORONAL SECTION PLANES

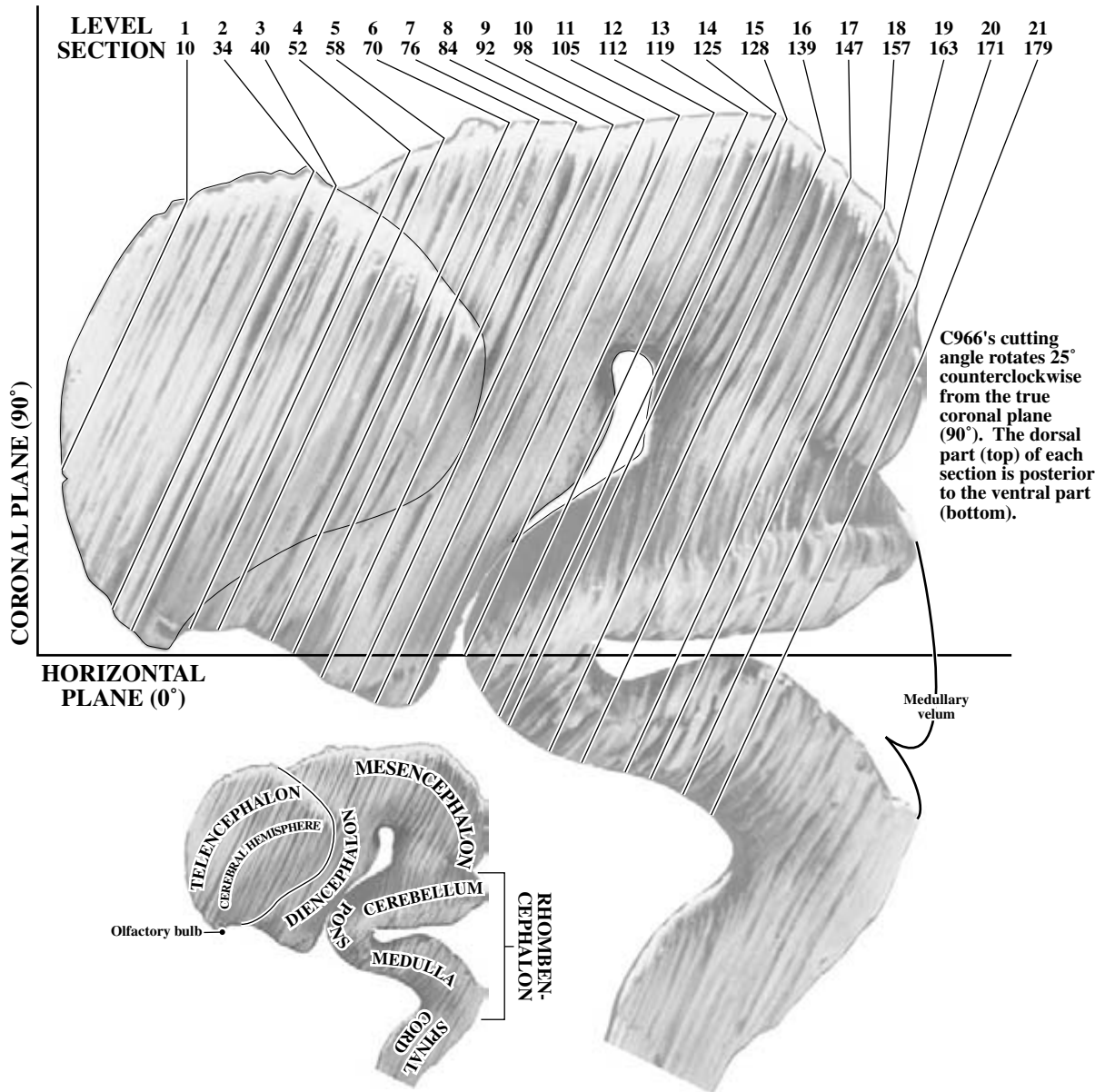


Figure 9. The lateral view of a 3-D model of C966's brain and upper cervical spinal cord (part of the Carnegie Collection at the National Museum of Health and Medicine) shows the exact locations and cutting angles of the illustrated sections of C966 in the following pages. The small inset identifies the major structural features. The medullary velum was not reconstructed so that the rhombencephalic superventricle appears as an open gap beneath the cerebellum.

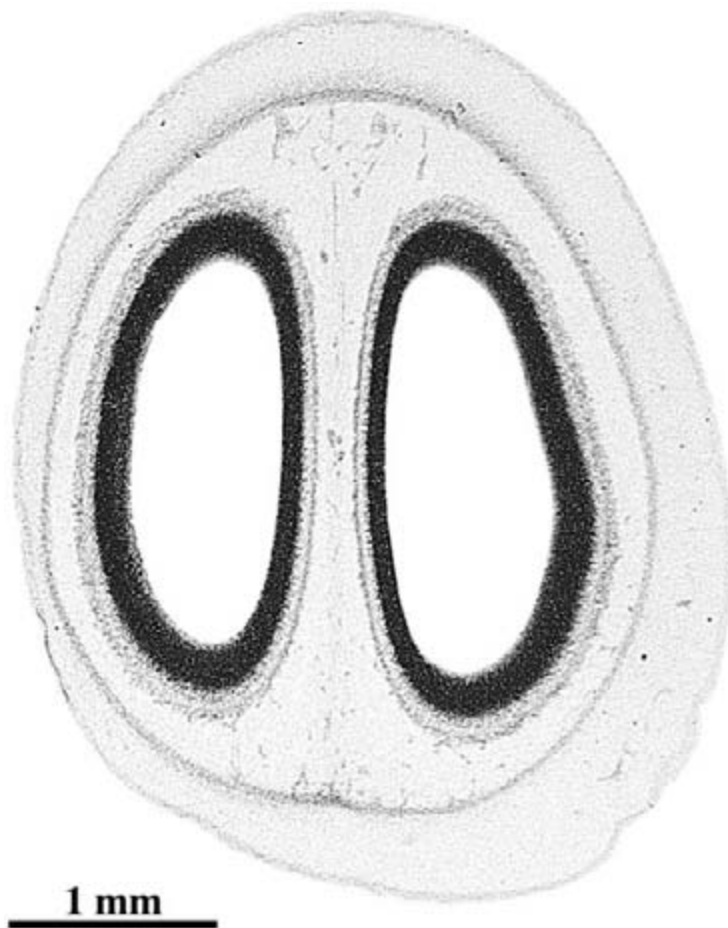
PLATE 186A

GW7.5 Coronal/horizontal

CR 23 mm

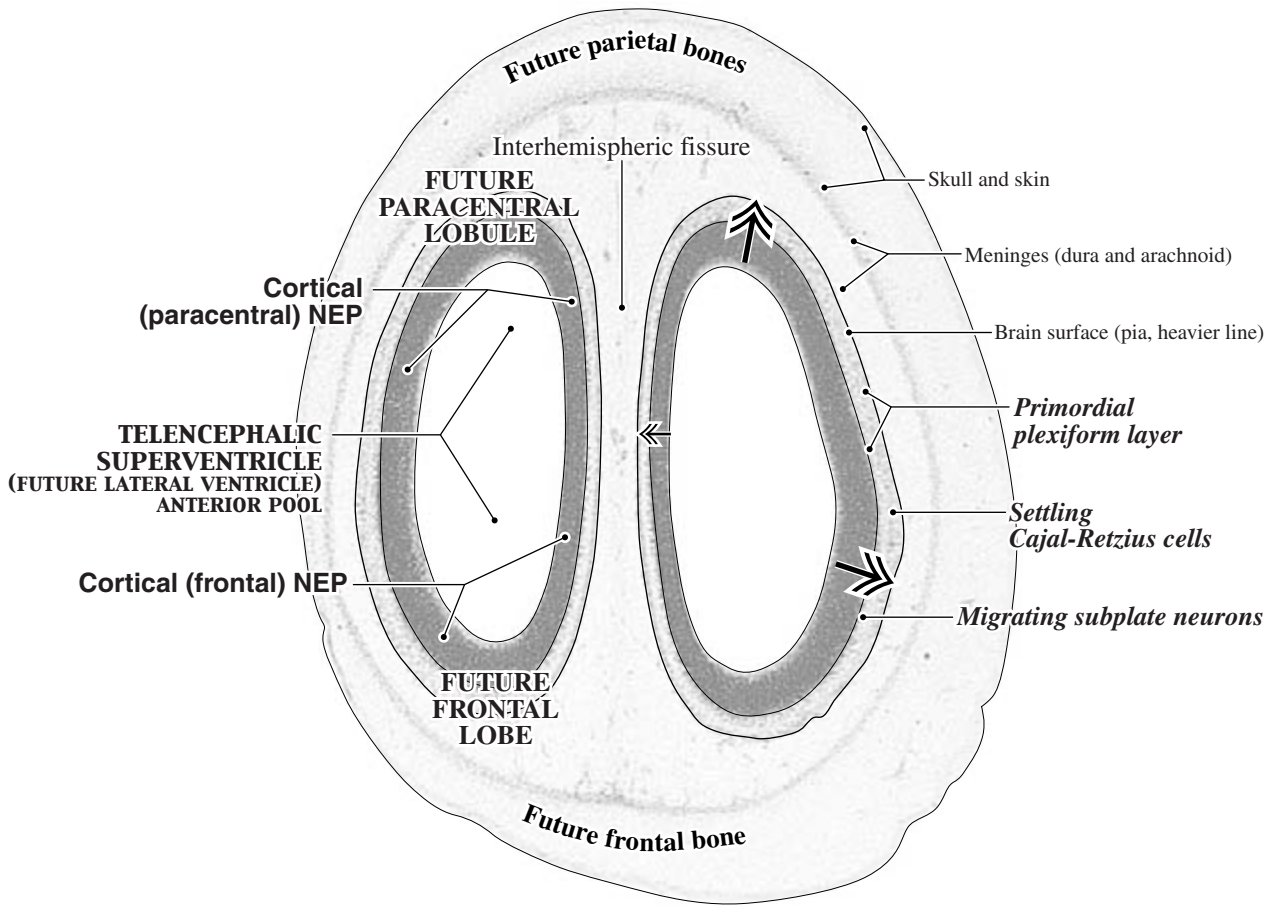
C966

Level 1: Section 10



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

NEP - Neuroepithelium



The cerebral cortex in this specimen does not have a *cortical plate* and a *stratified transitional field* outside the cortical neuroepithelium. Instead, there is a *primordial plexiform layer* composed of early-generated Cajal-Retzius cells and the slightly later-generated subplate neurons.

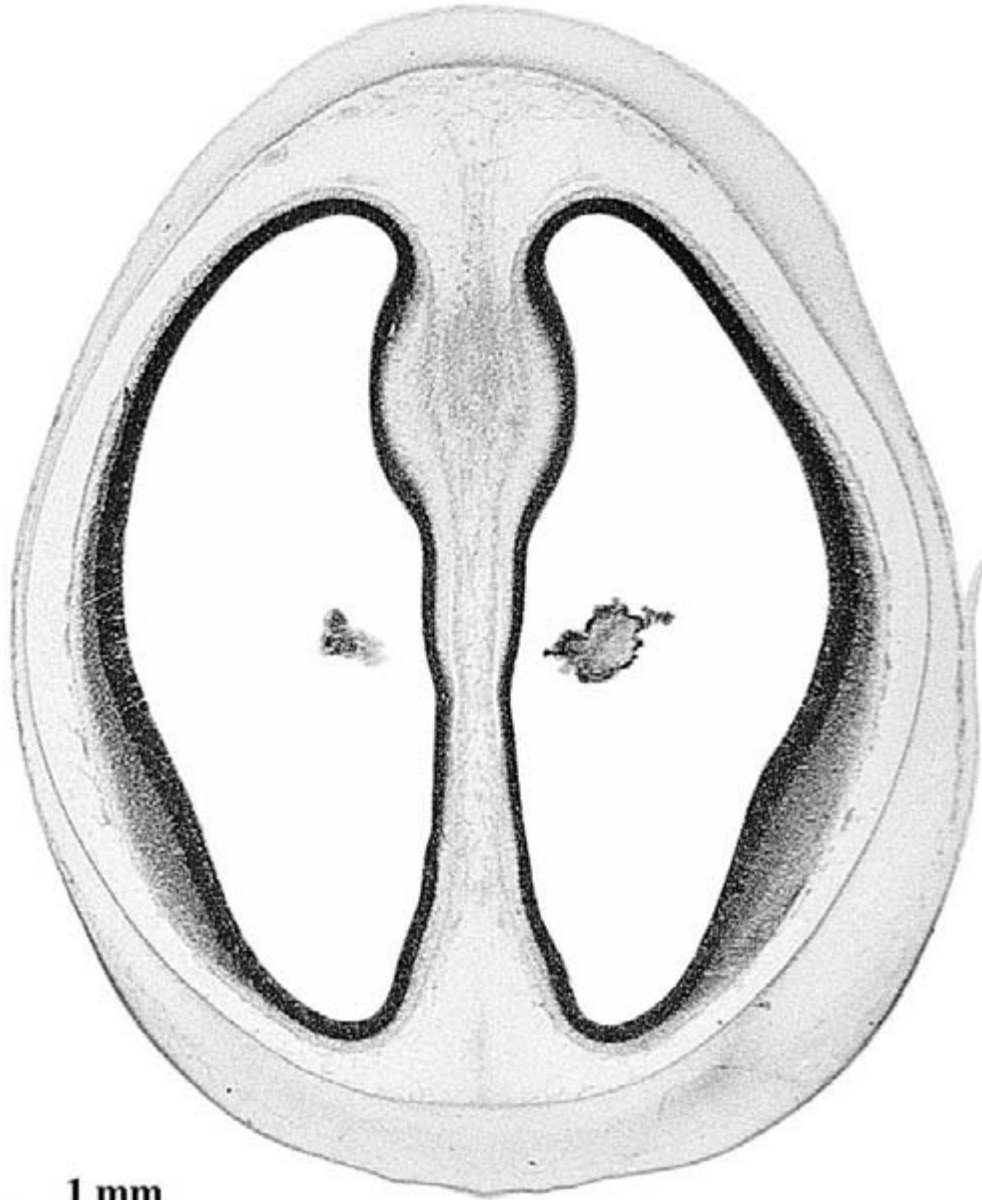
The large Cajal-Retzius cells settle subjacent to the pia meninx and remain superficial in cortical Layer I throughout later development. The deep border of Layer I is not defined until a cortical plate appears.

Subplate neurons accumulate in a loosely defined network beneath the Cajal-Retzius cells. In later development they become radially aligned and are the pioneer neurons in the cortical plate. Eventually, they delaminate and settle in the subplate (cortical Layer VII).

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 187A

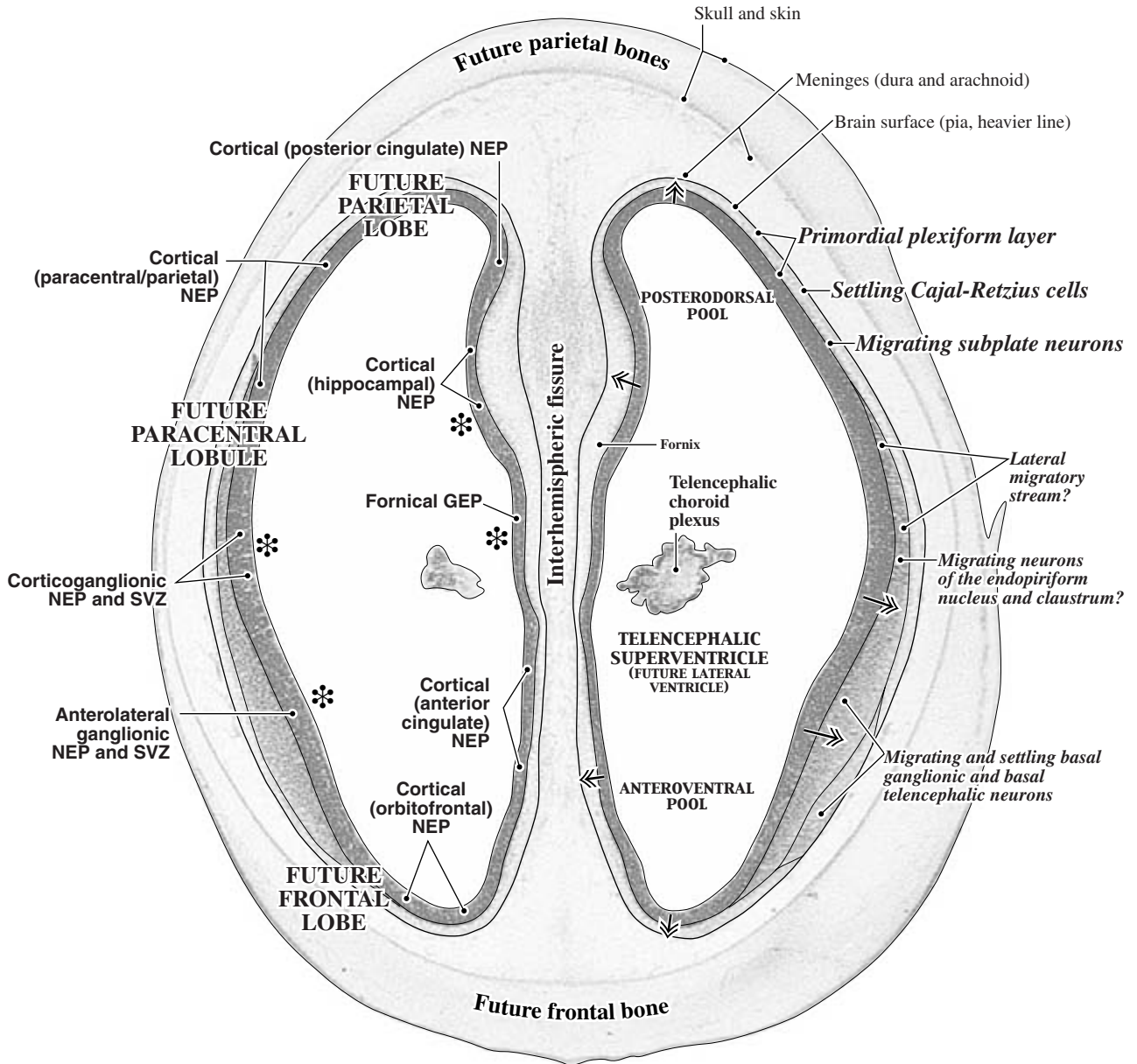
**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 2: Section 34**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

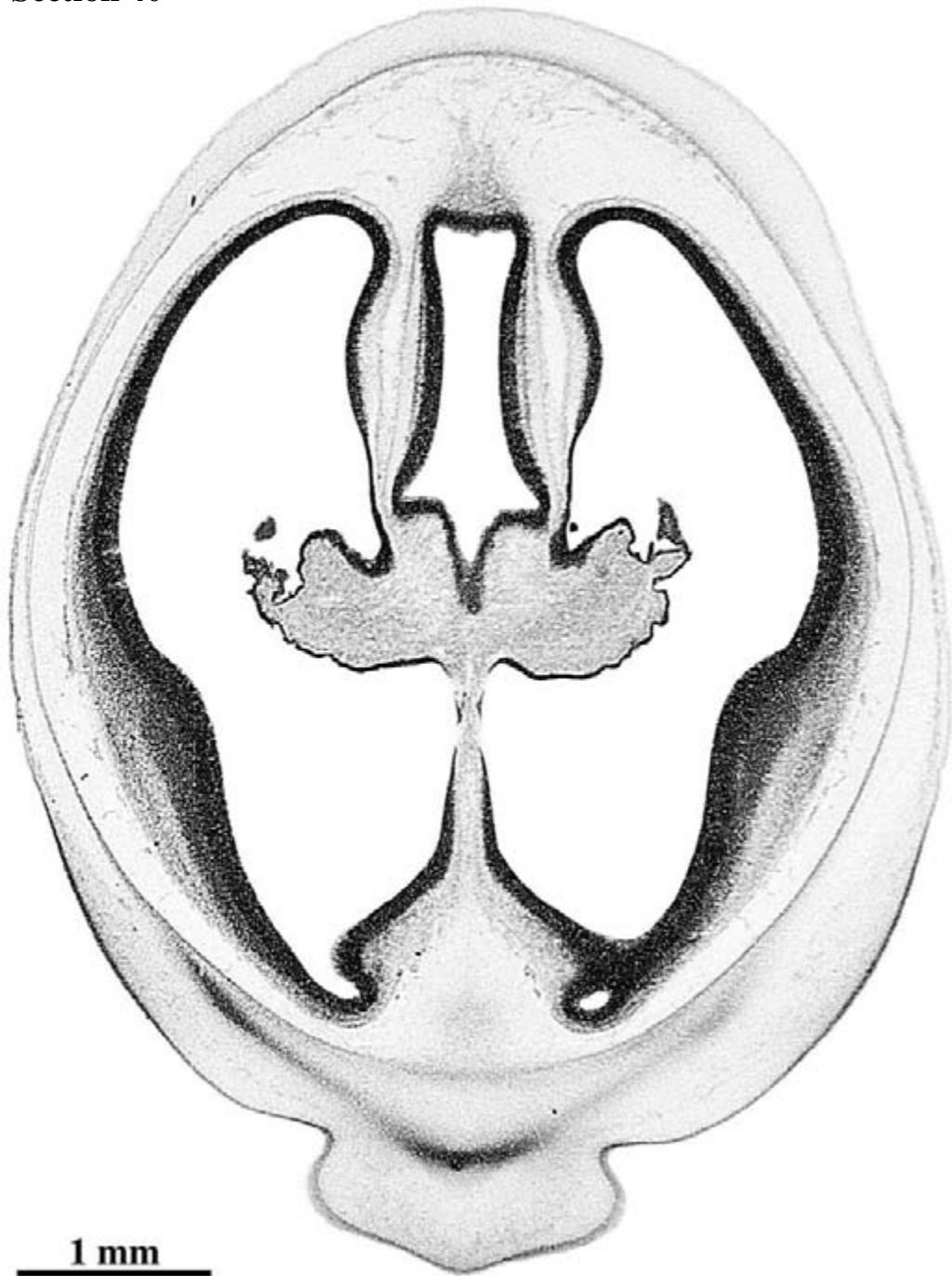


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

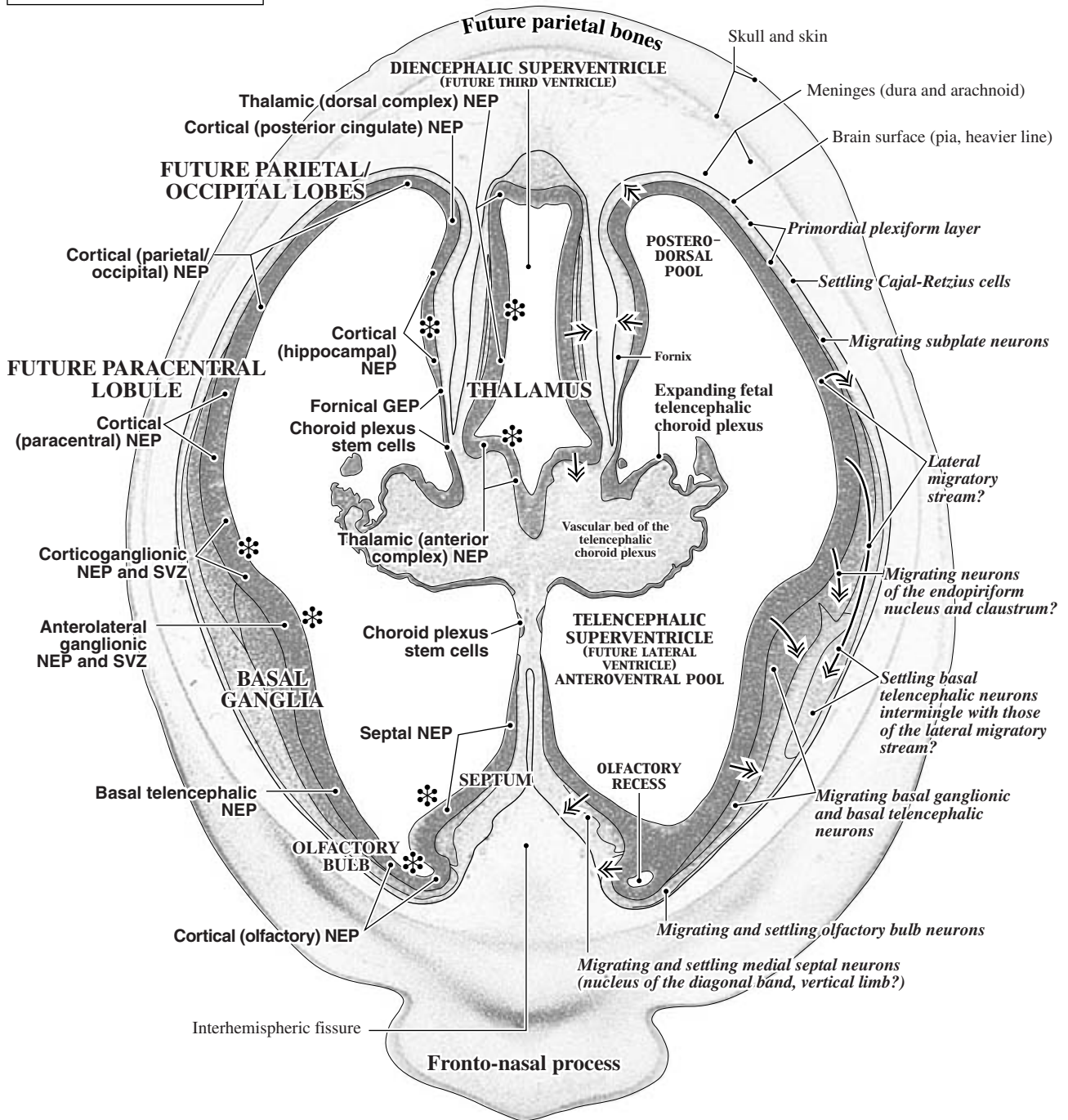
PLATE 188A

**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 3: Section 40**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

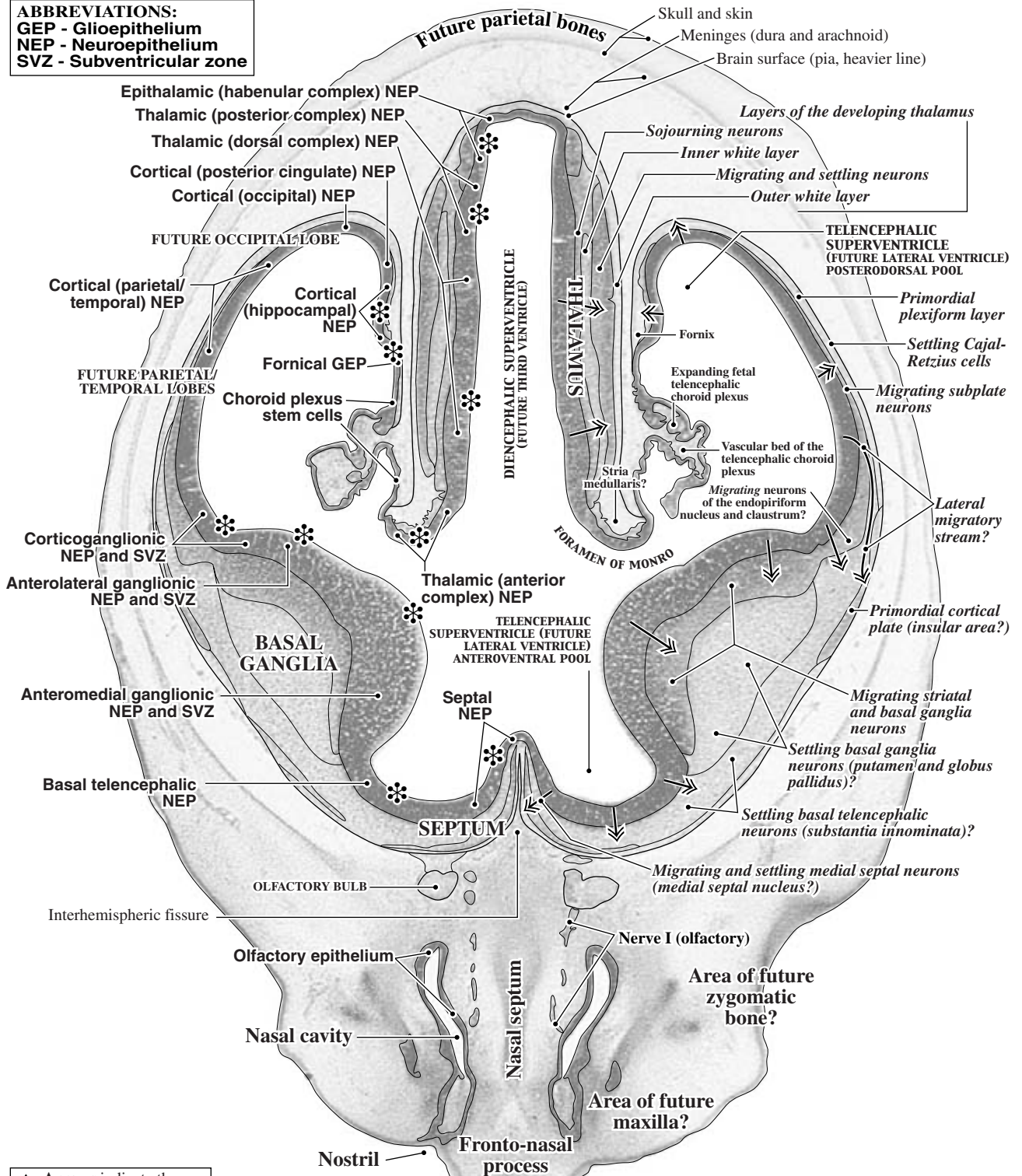
PLATE 189A

**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 4: Section 52**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

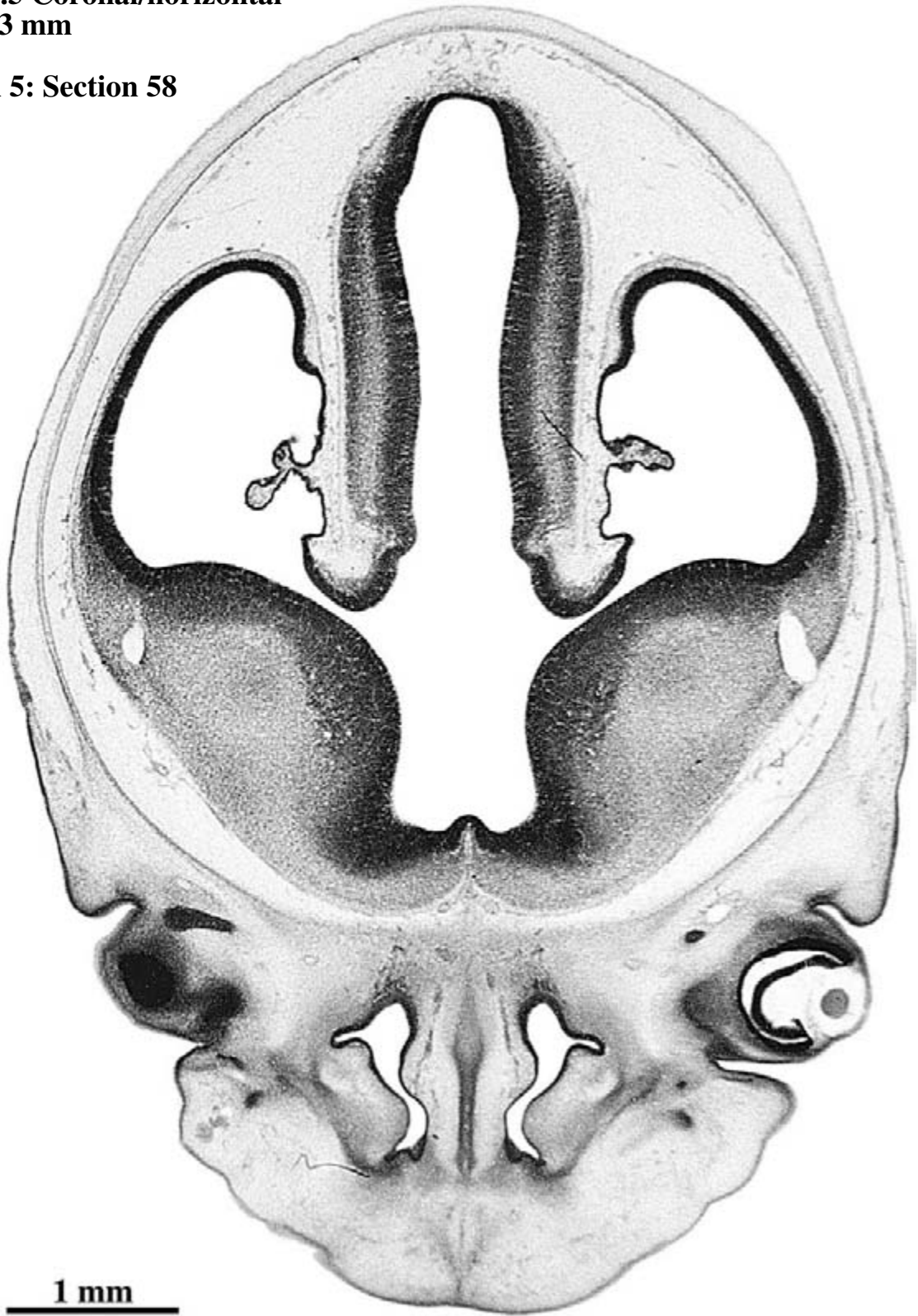


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

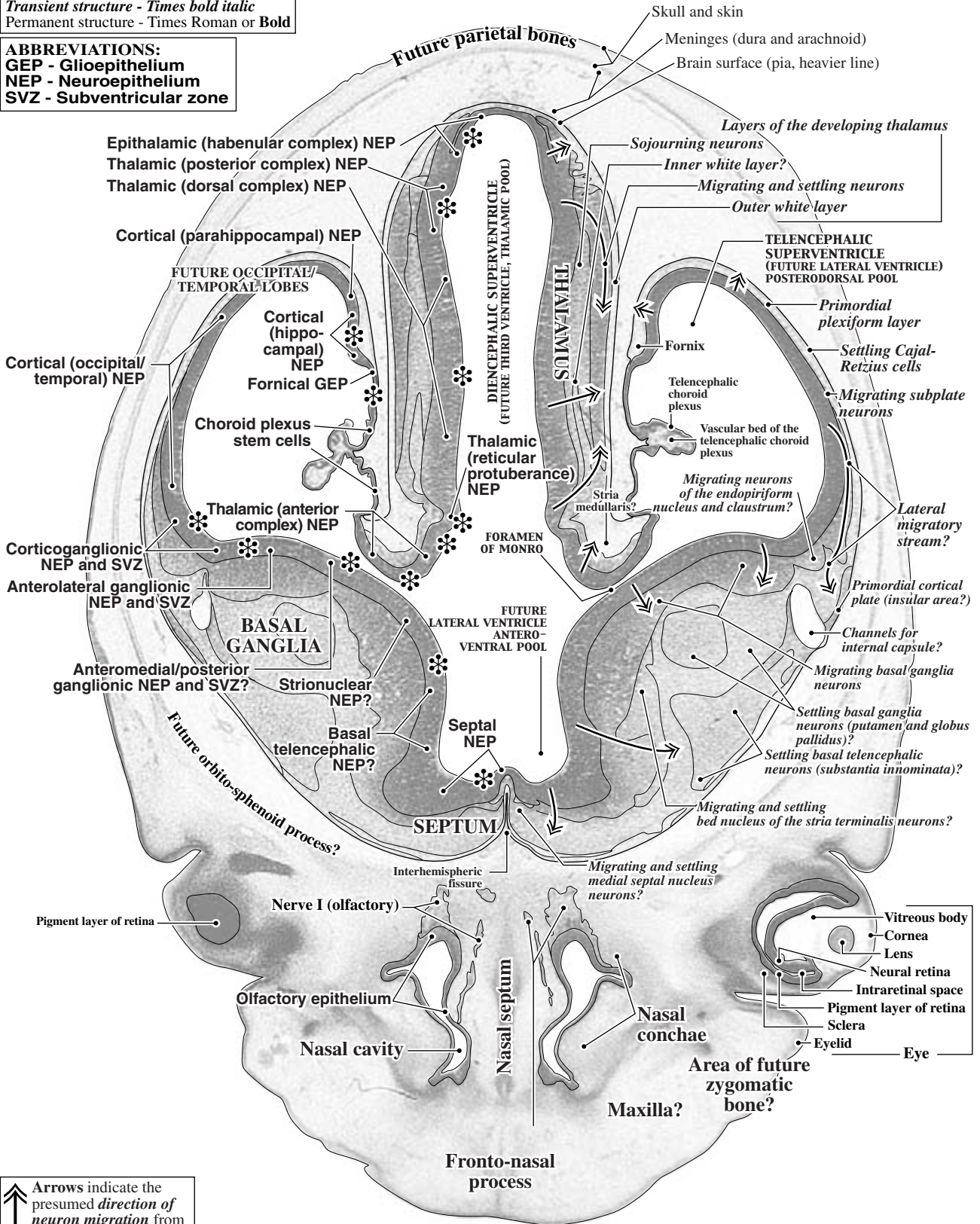
PLATE 190A

**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 5: Section 58**



FONT KEY:
VENTRICULAR DIVISIONS – CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 191A

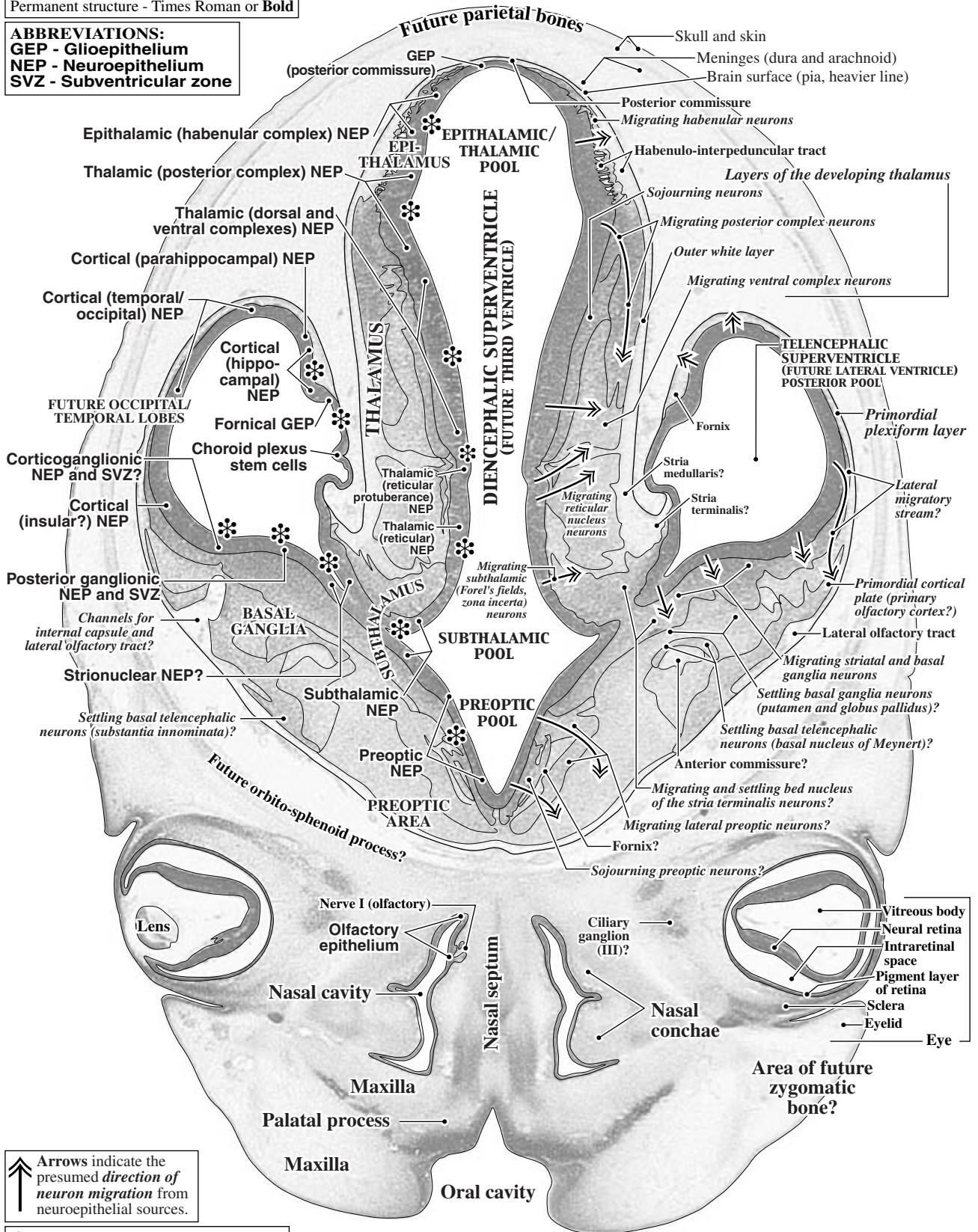
**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 6: Section 70**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

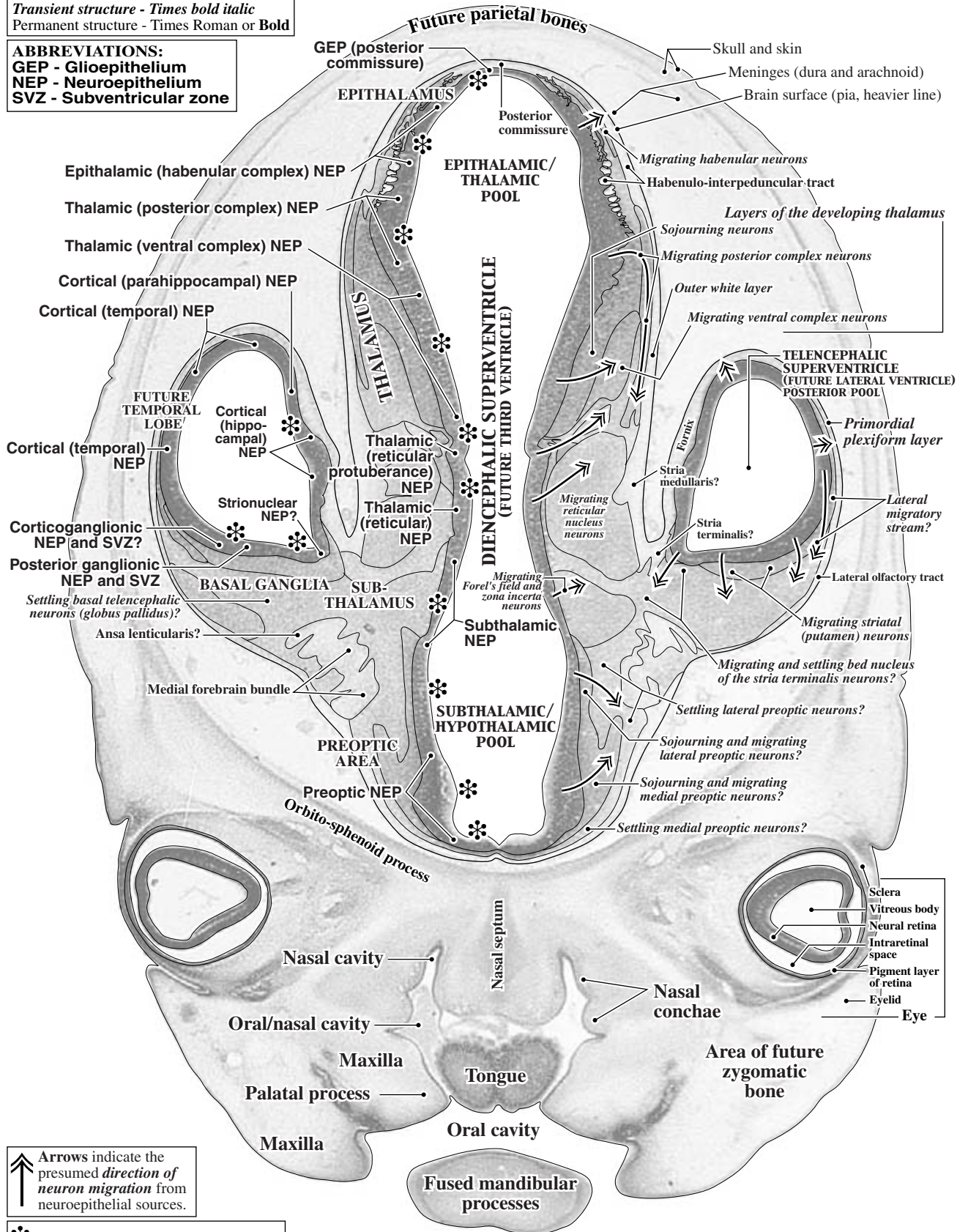
PLATE 192A

**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 7: Section 76**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone

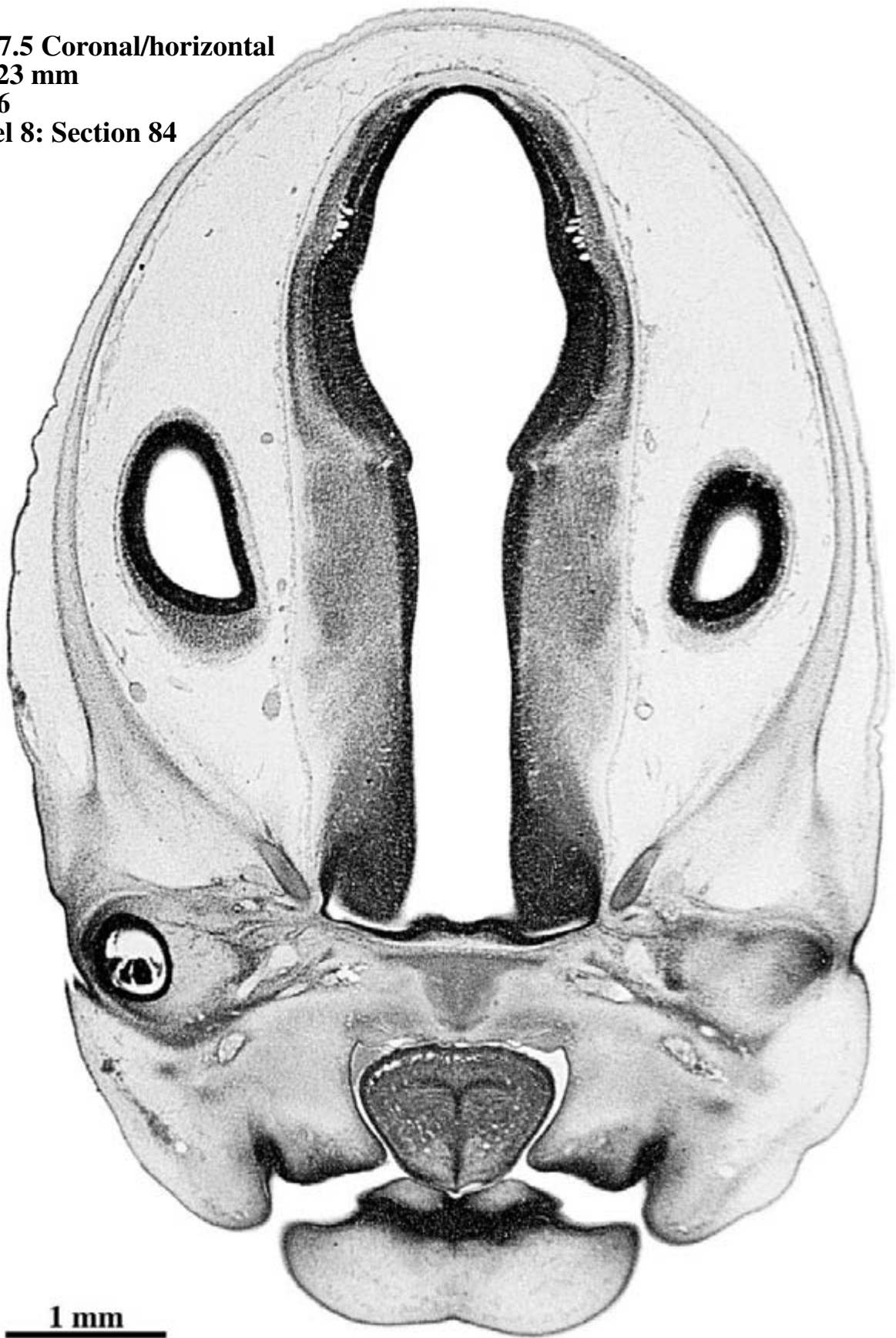


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 193A

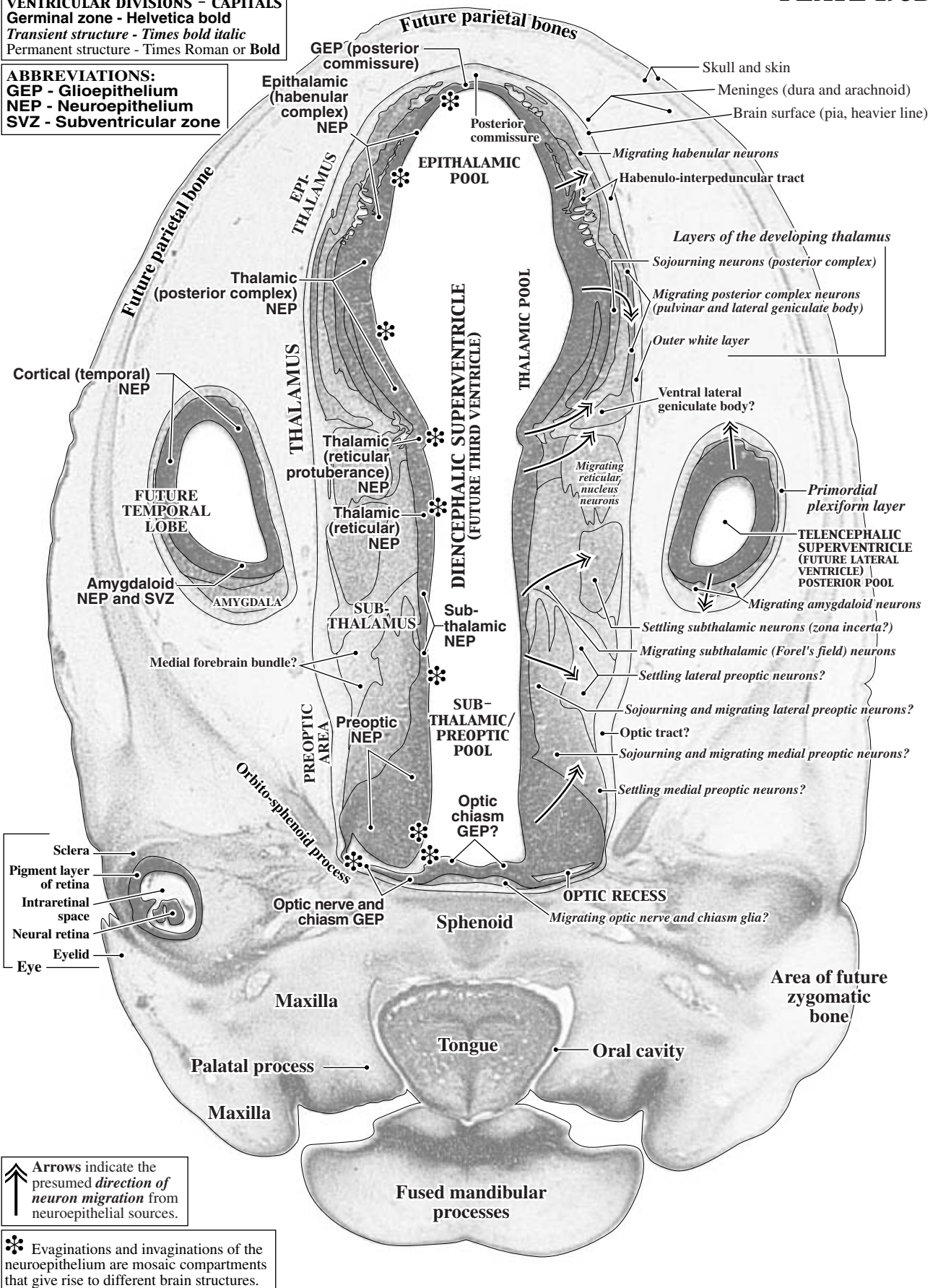
**GW7.5 Coronal/horizontal
CR 23 mm
C966
Level 8: Section 84**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
SVZ - Subventricular zone



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

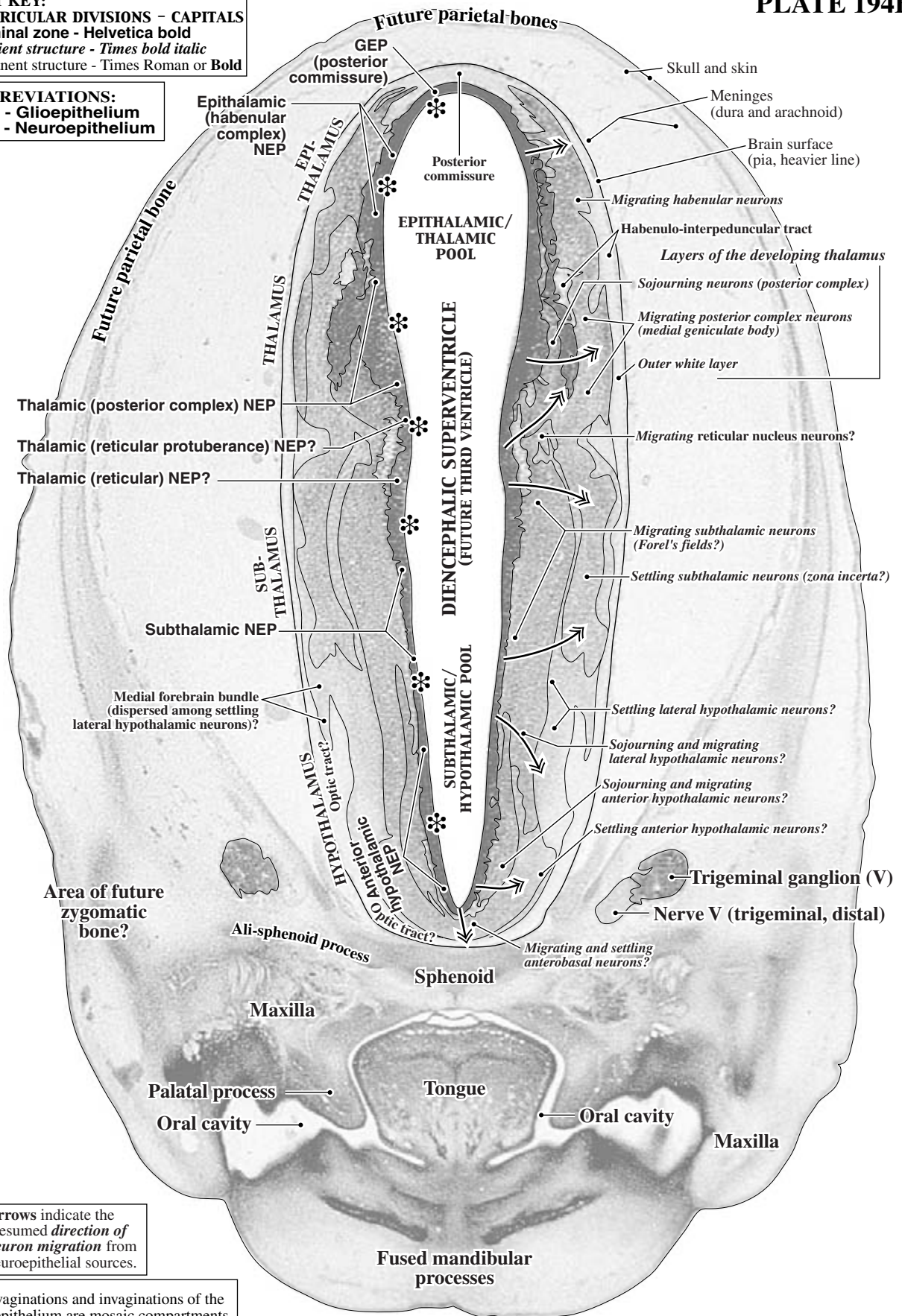
PLATE 194A

**GW7.5
Coronal/horizontal
CR 23 mm
C966
Level 9:
Section 92**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

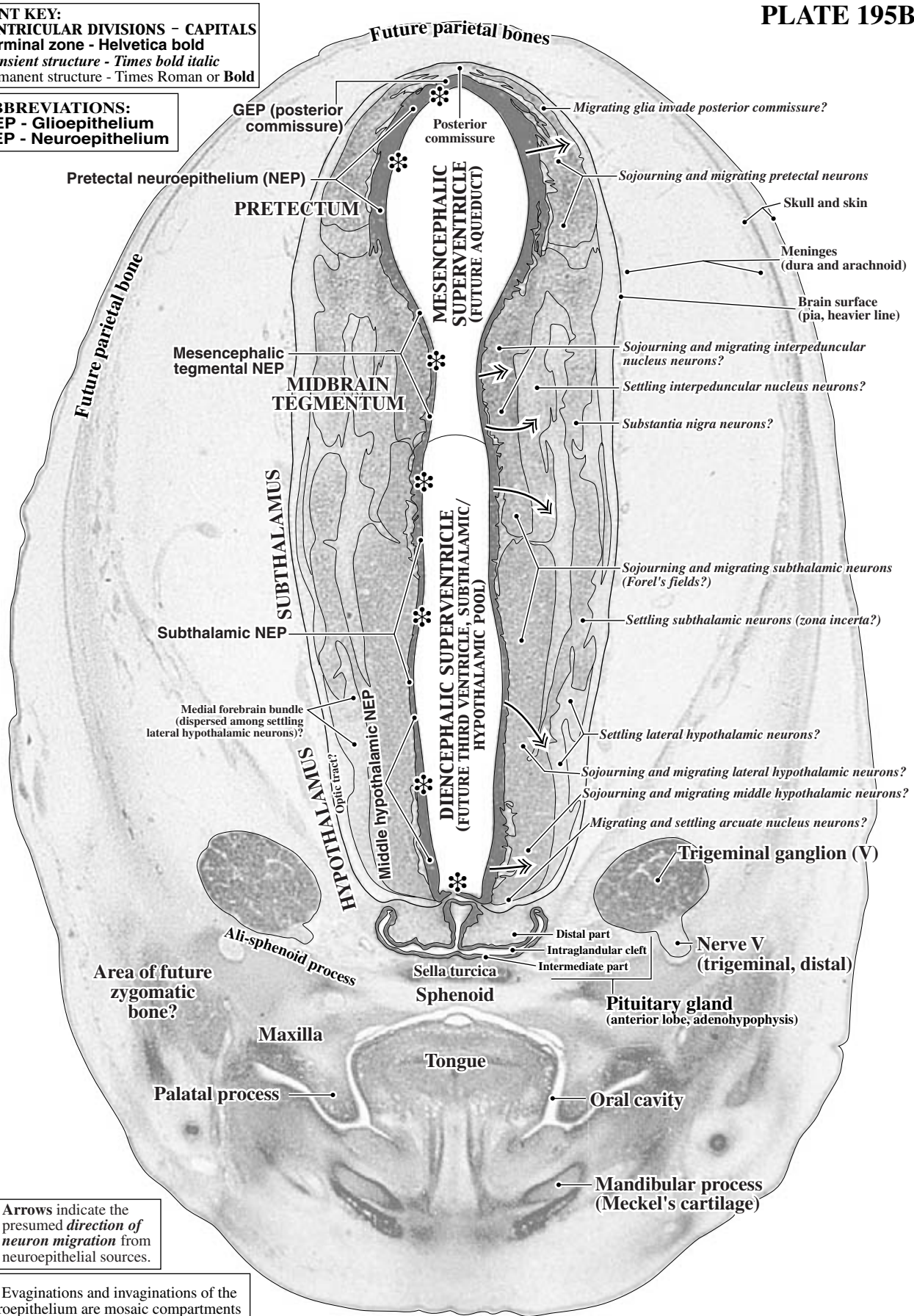
PLATE 195A

GW7.5
Coronal/horizontal
CR 23 mm, C966
Level 10:
Section 98



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 196A

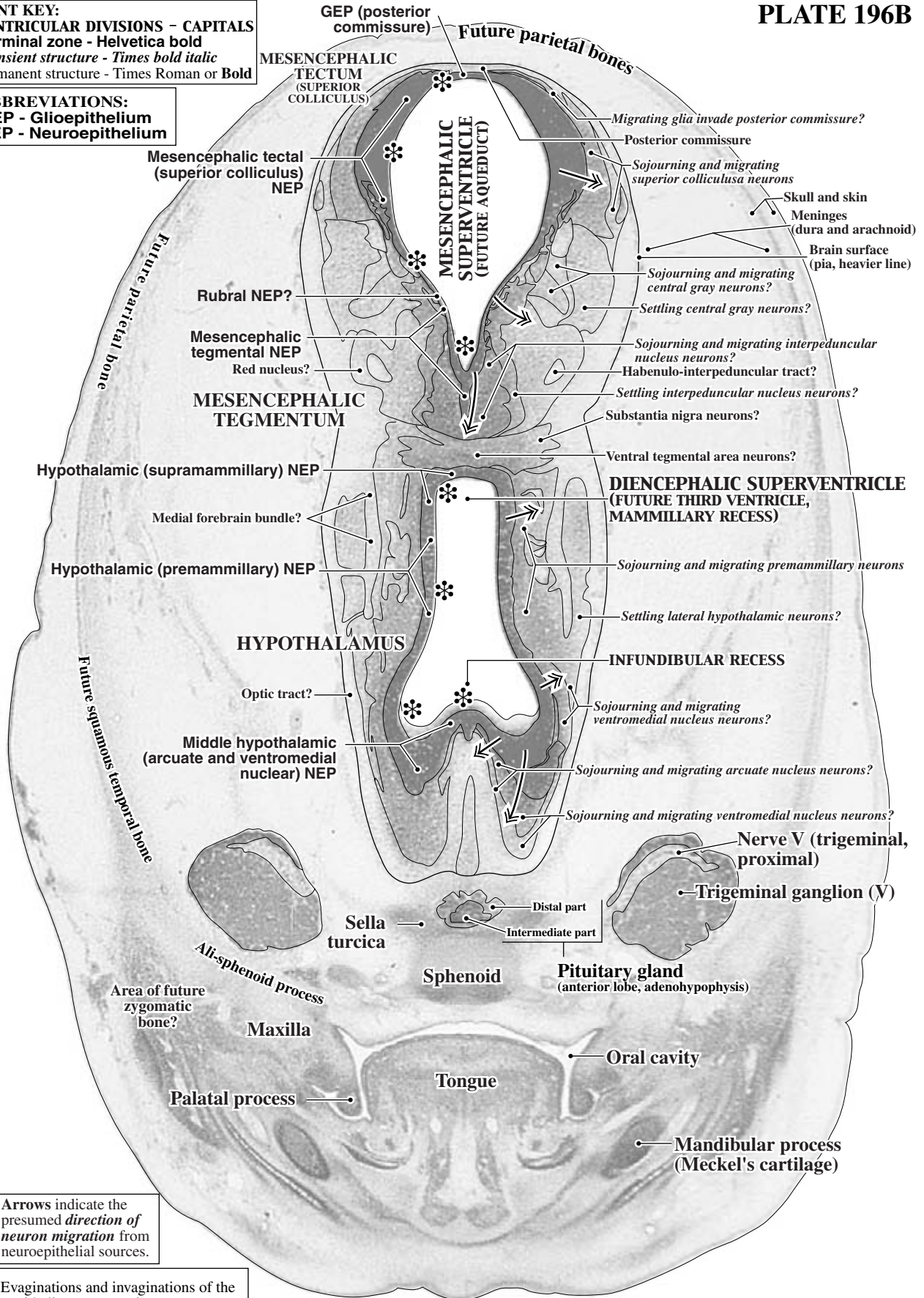
GW7.5
Coronal/horizontal
CR 23 mm, C966
Level 11:
Section 105



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

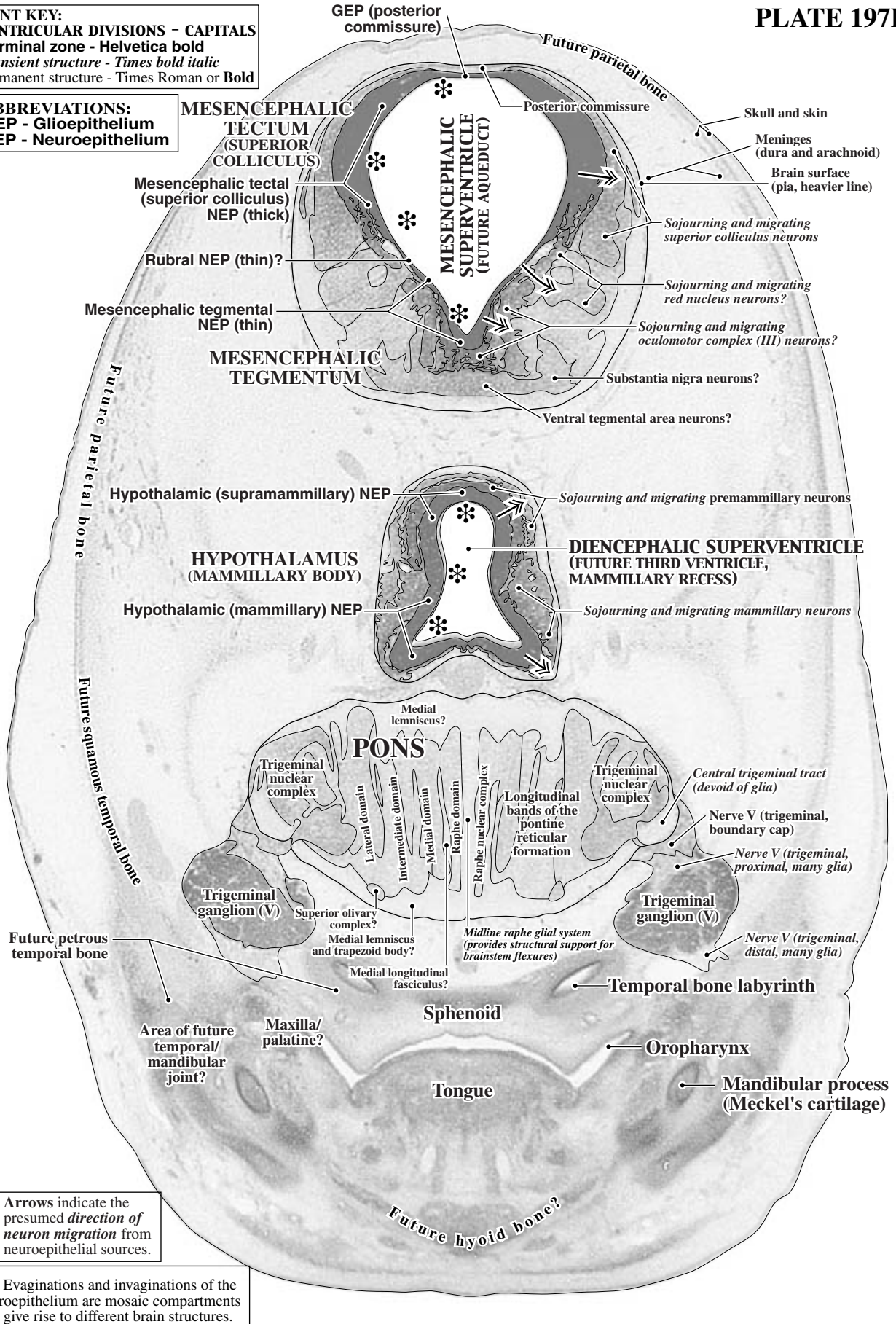
PLATE 197A

GW7.5
Coronal/horizontal
CR 23 mm, C966
Level 12:
Section 112



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 198A

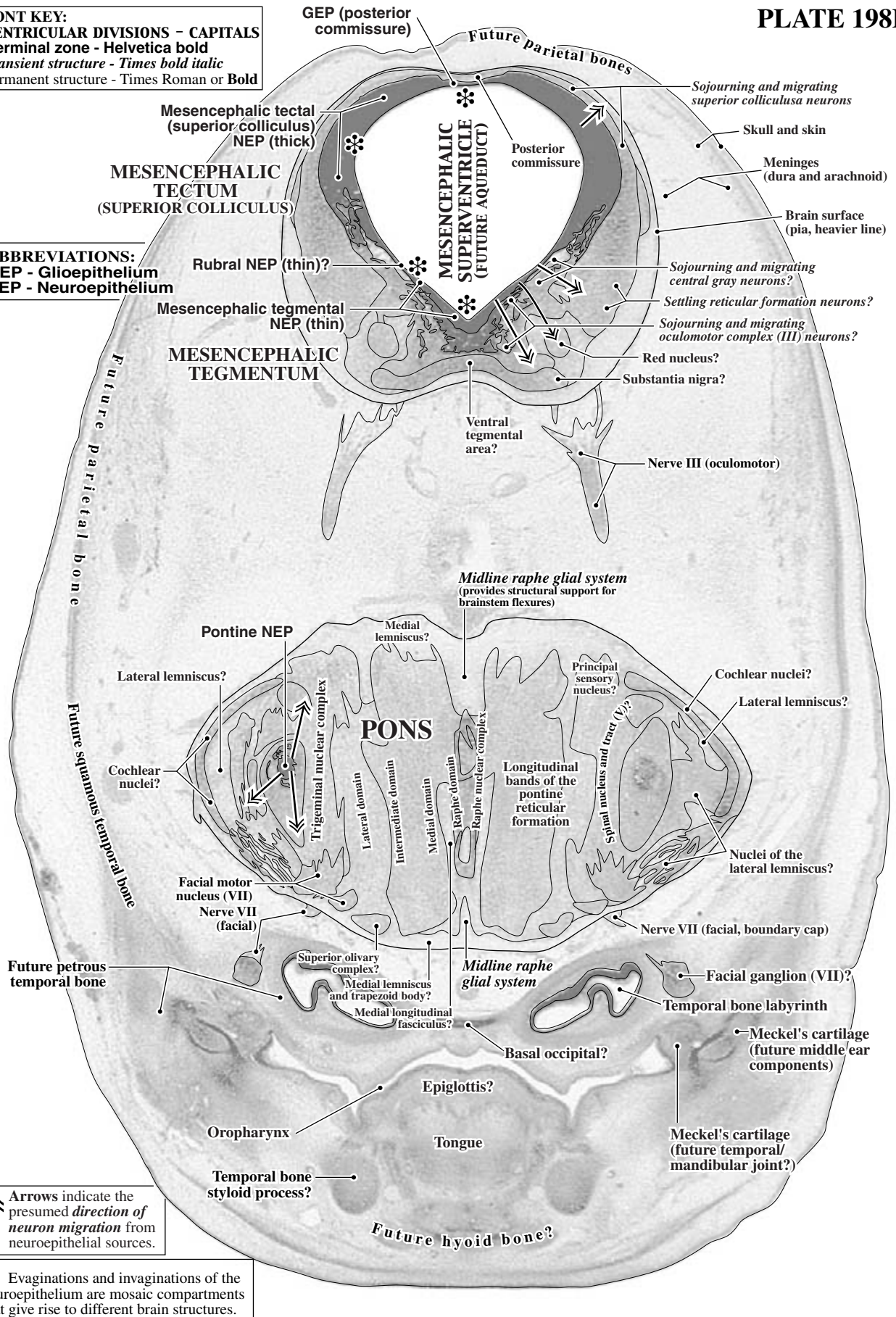
GW7.5
Coronal/horizontal
CR 23 mm, C966
Level 13:
Section 119



PLATE 198B

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium

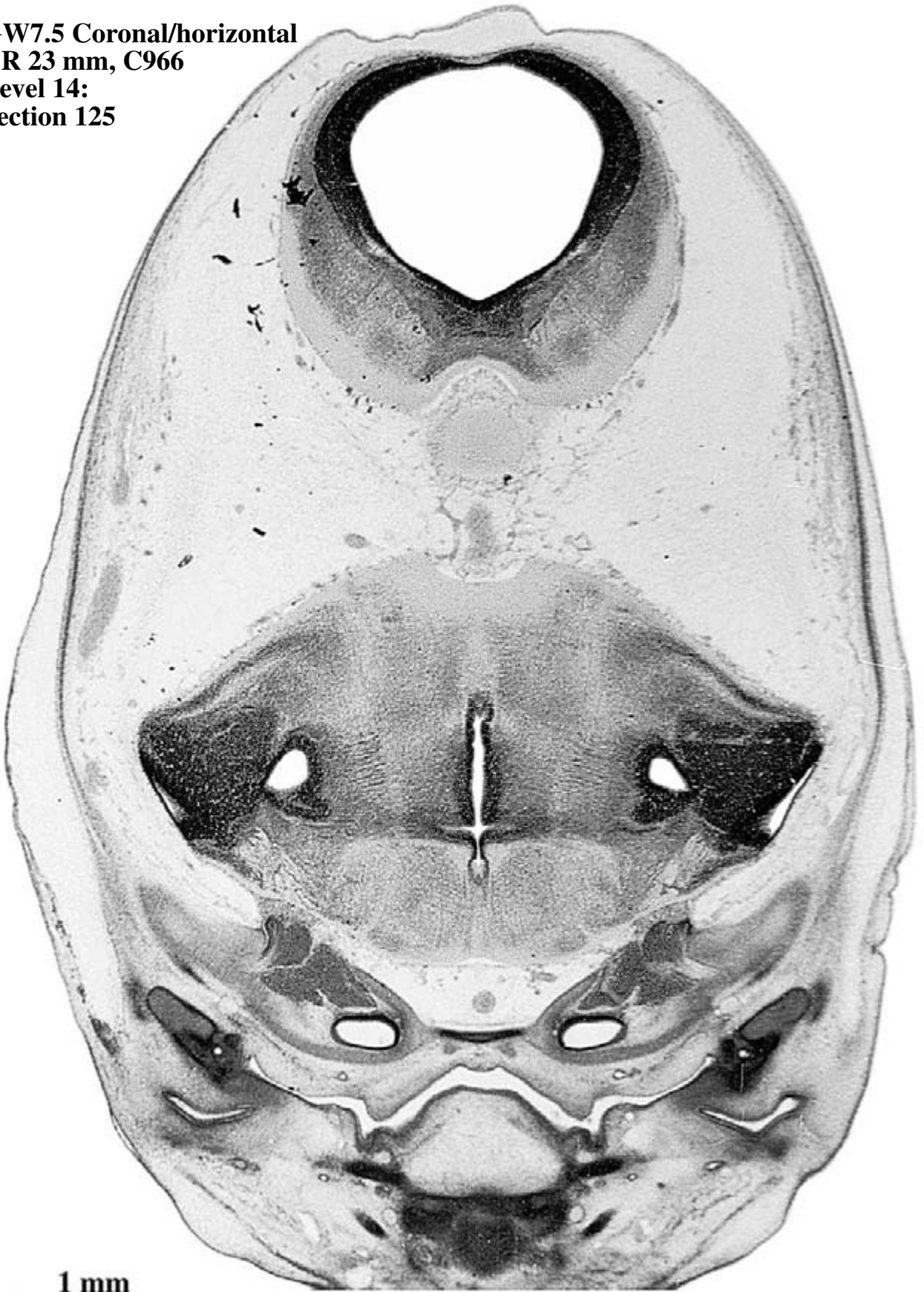


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 199A

**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 14:
Section 125**

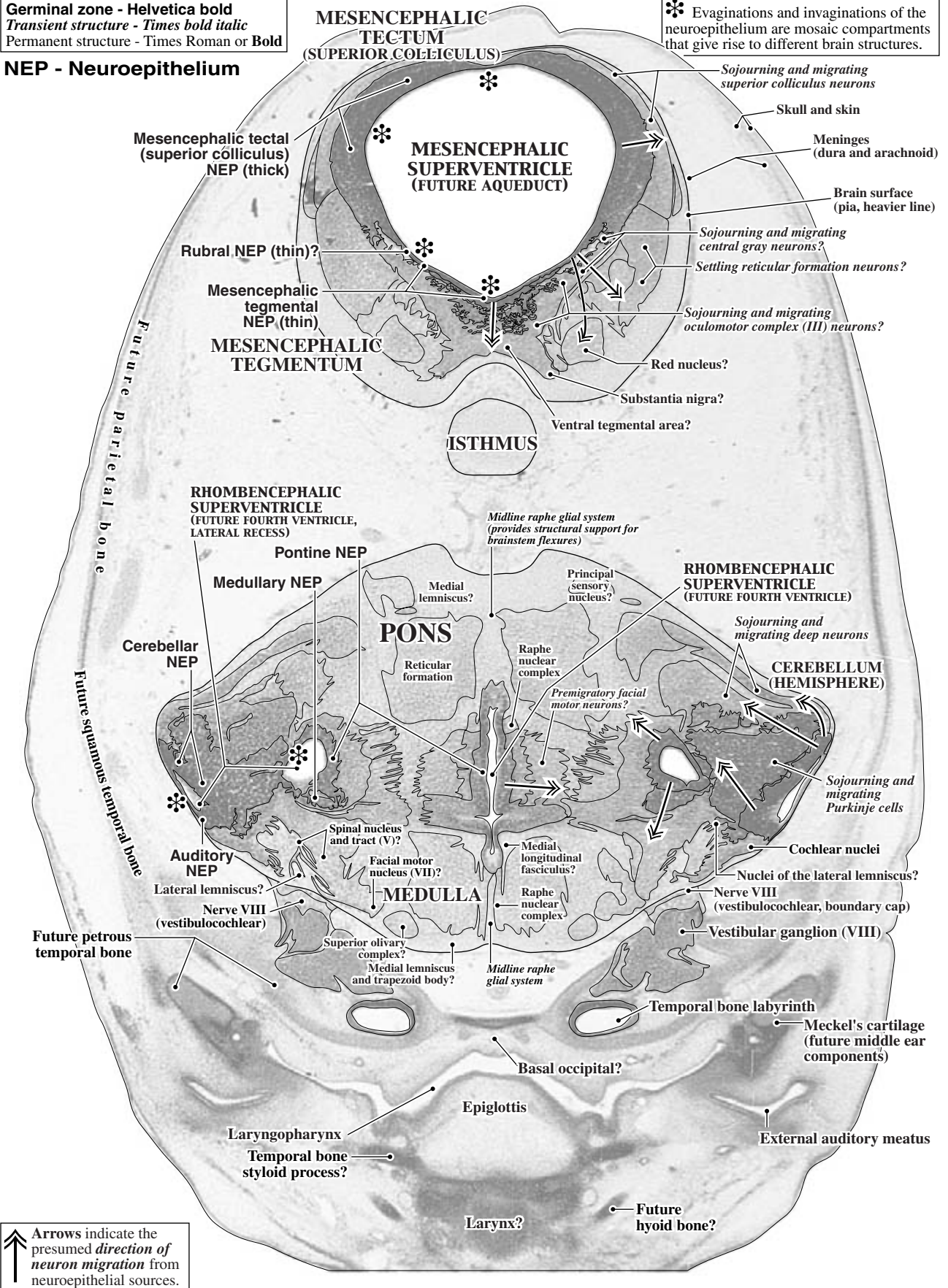


1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

NEP - Neuroepithelium

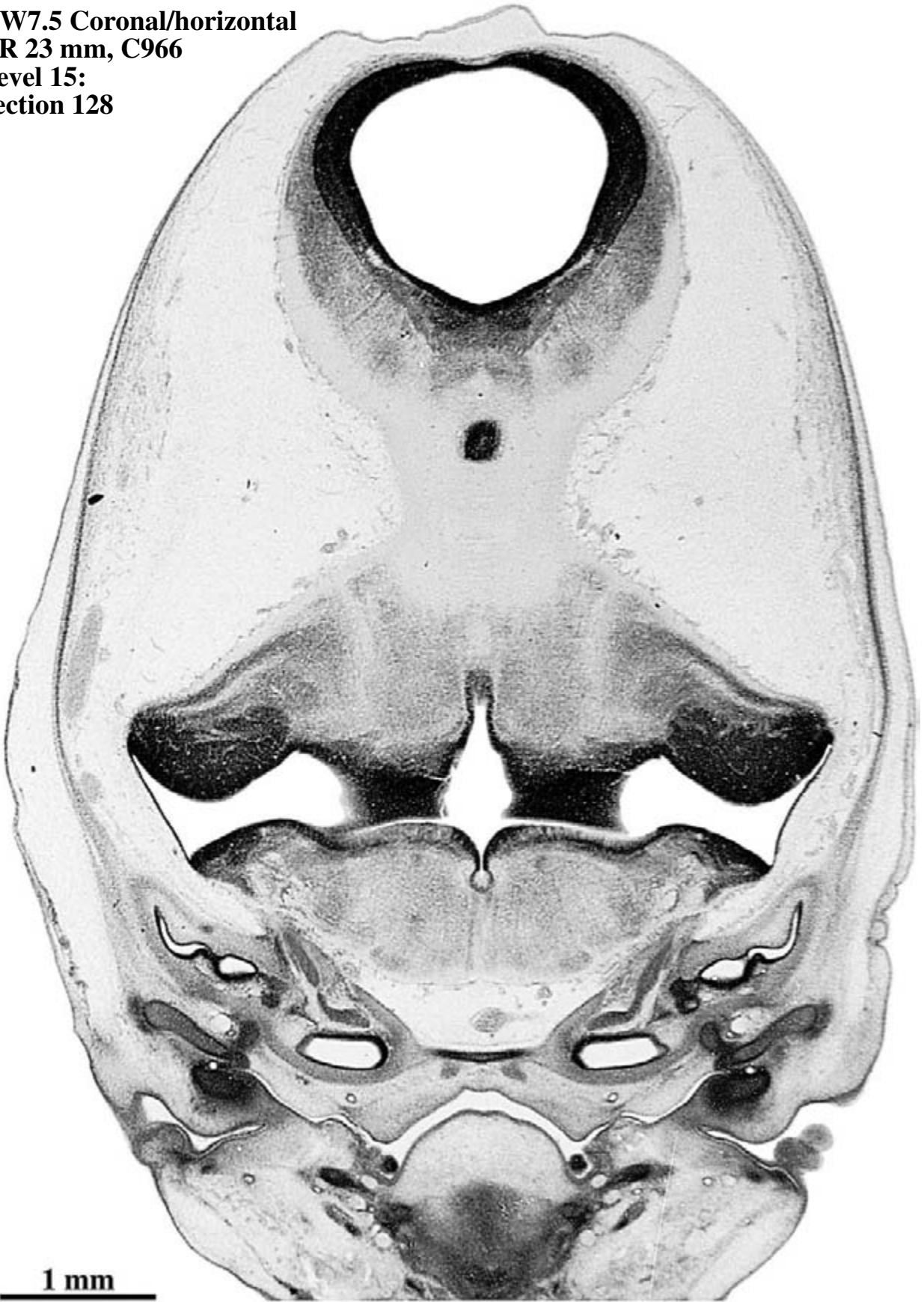
* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 200A

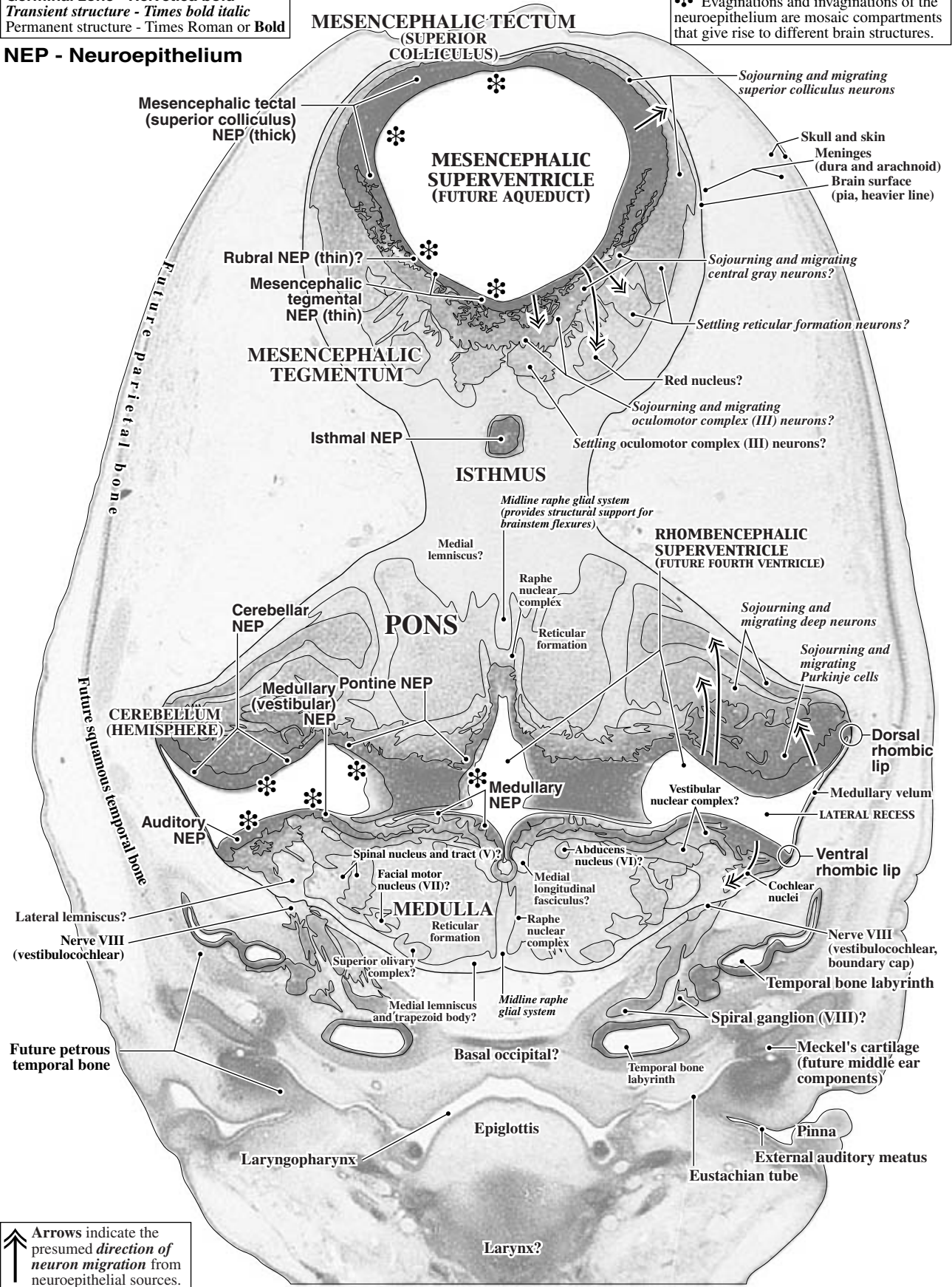
**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 15:
Section 128**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

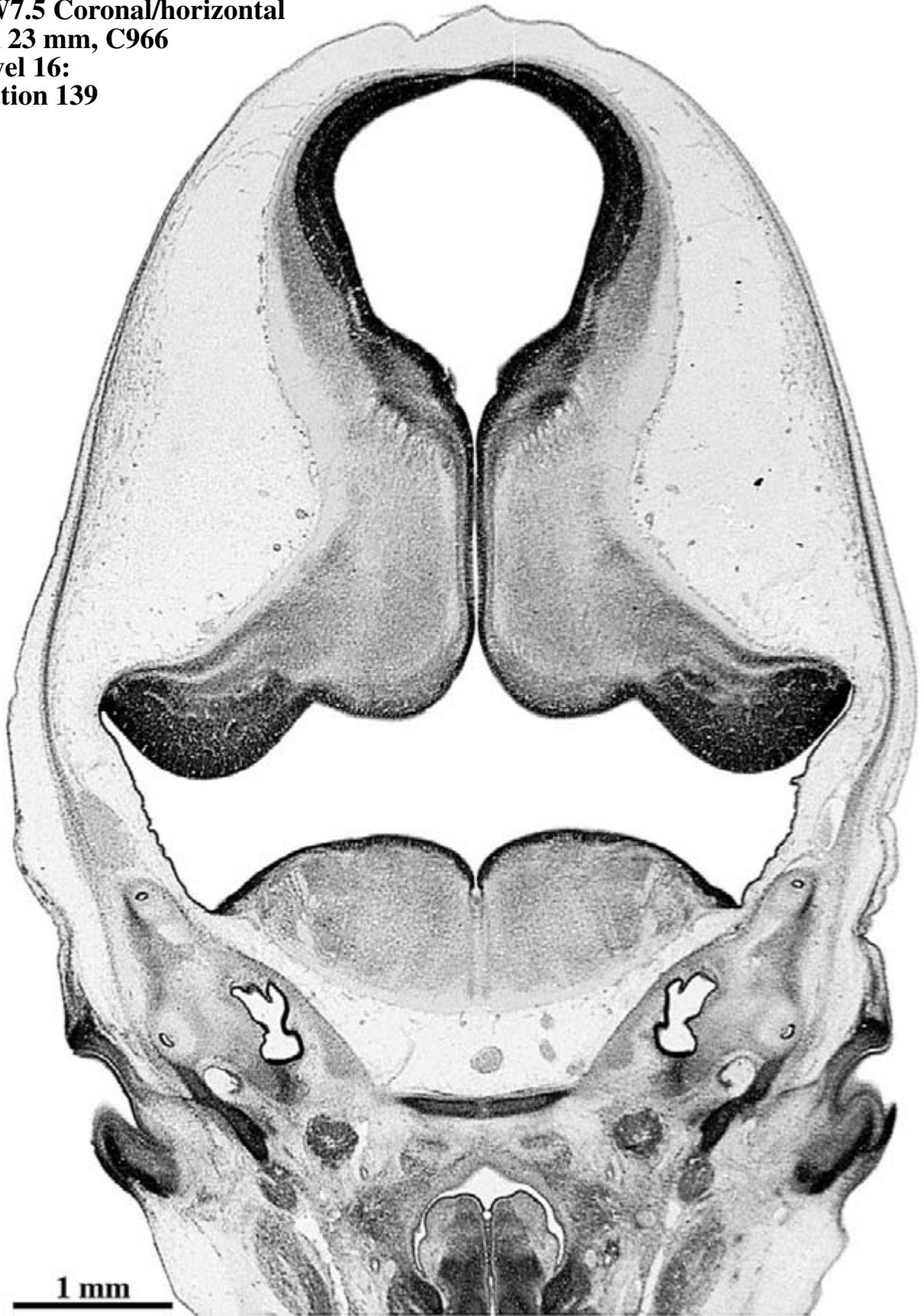
NEP - Neuroepithelium



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

PLATE 201A

**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 16:
Section 139**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
NEP - Neuroepithelium
CTF - Cerebellar transitional field

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

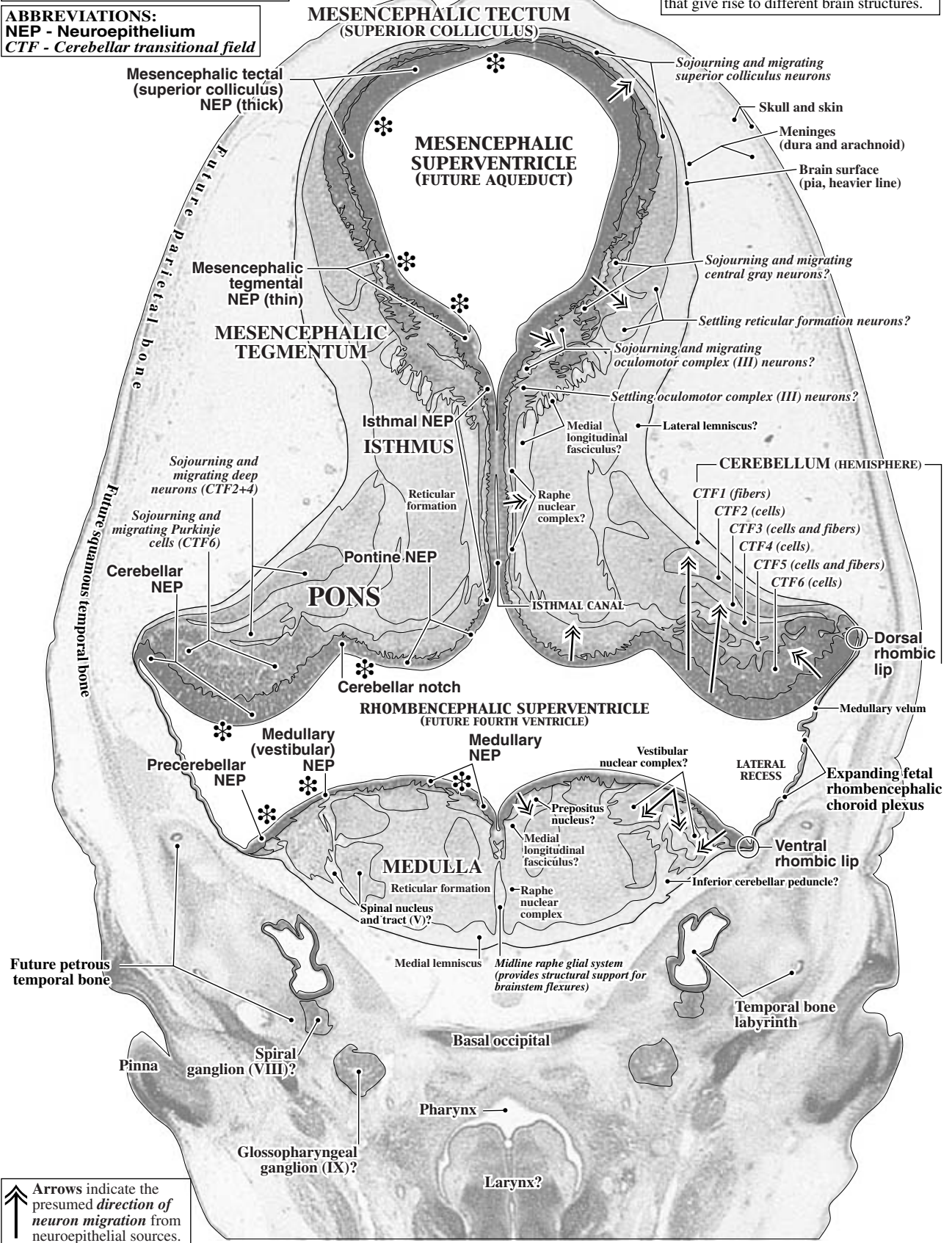
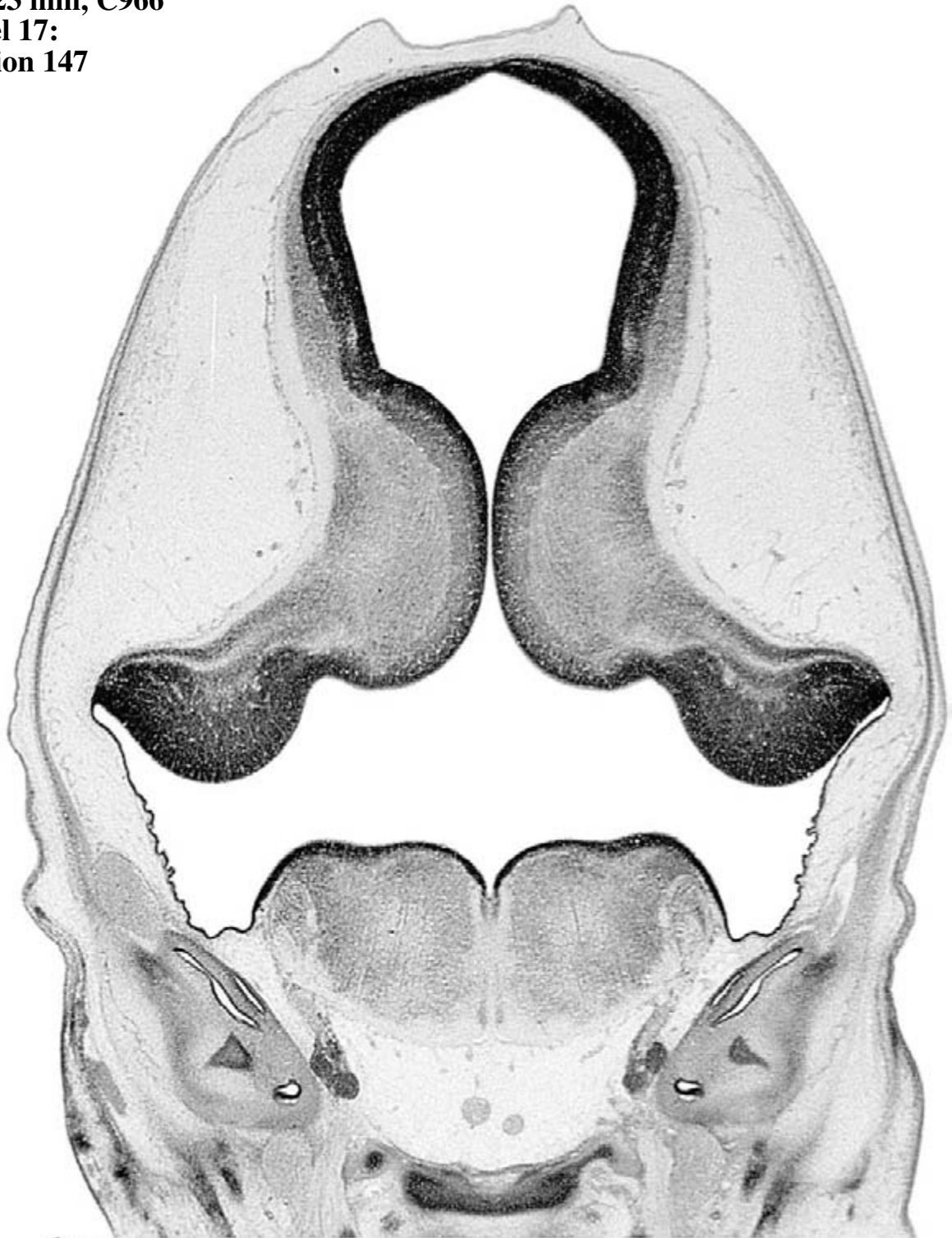


PLATE 202A

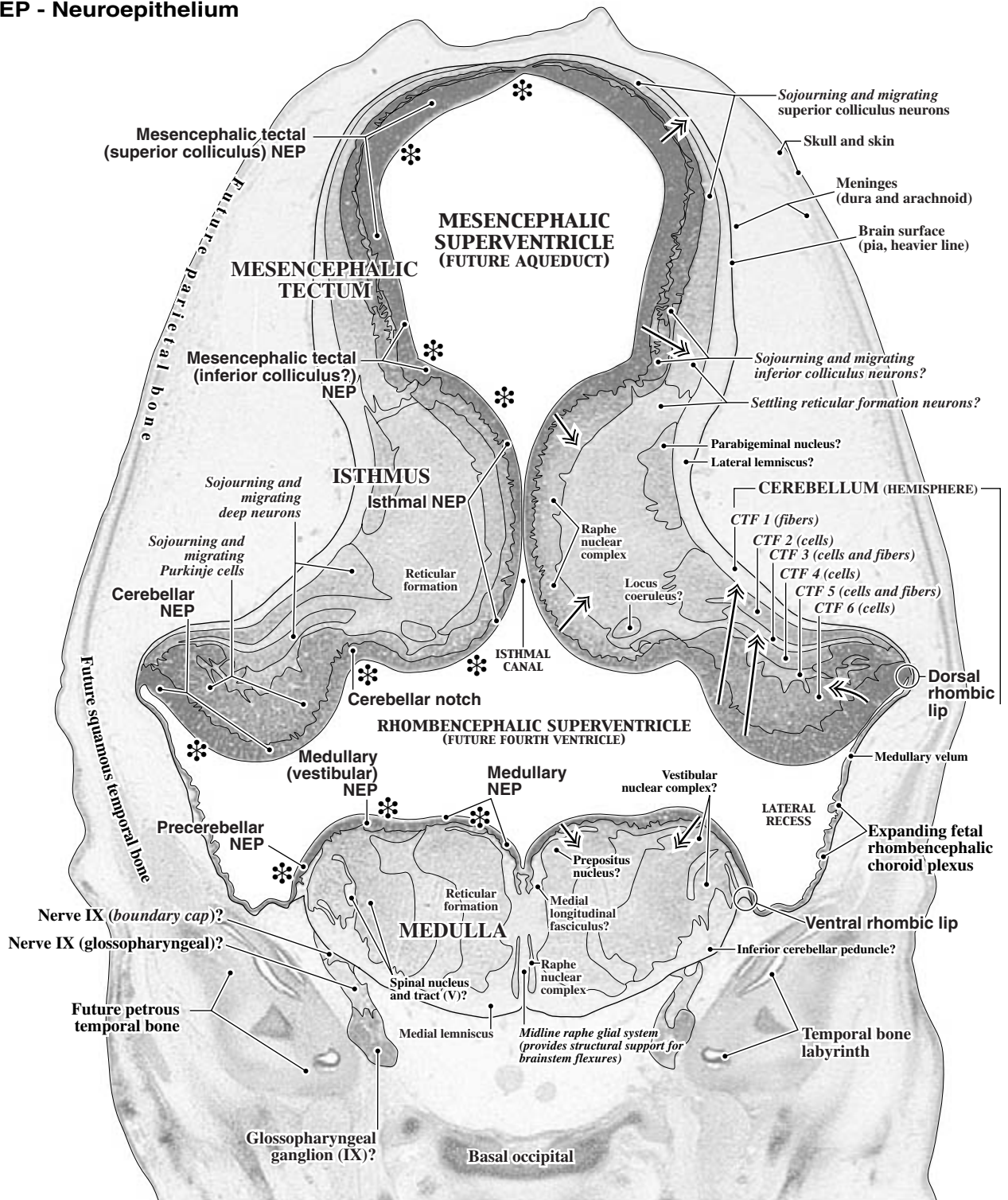
**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 17:
Section 147**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

NEP - Neuroepithelium

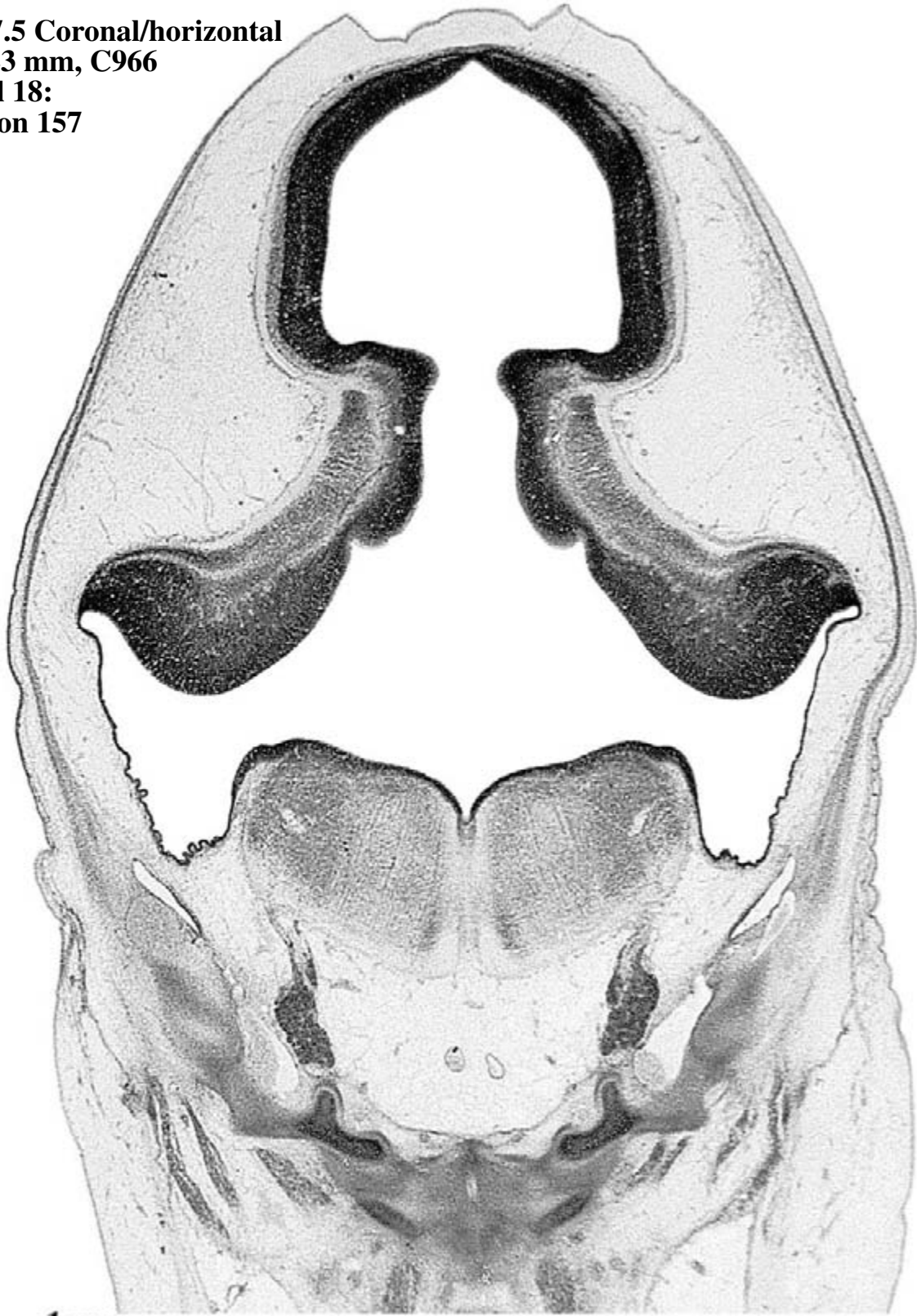


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 203A

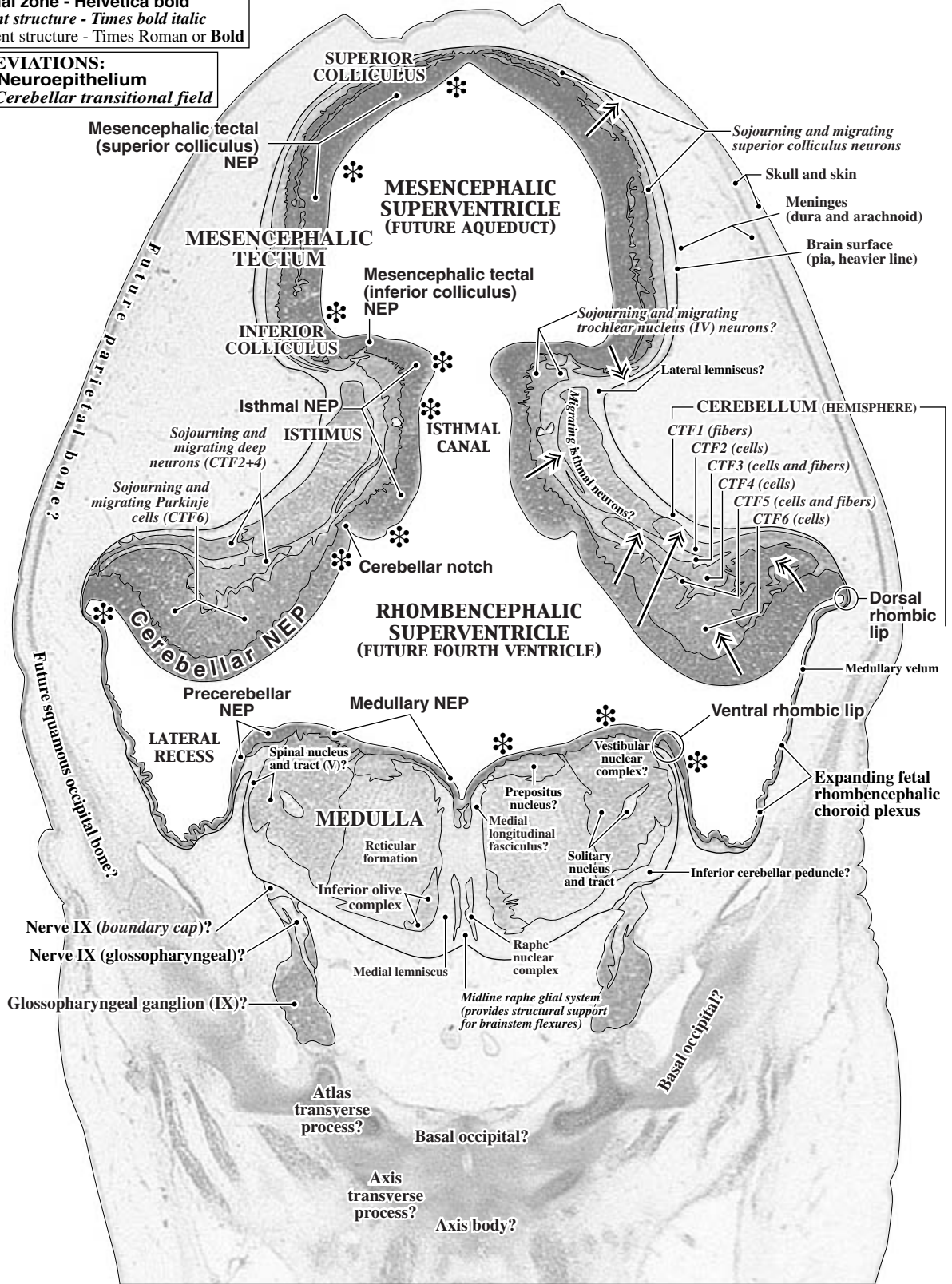
**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 18:
Section 157**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
NEP - Neuroepithelium
CTF - Cerebellar transitional field



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 204A

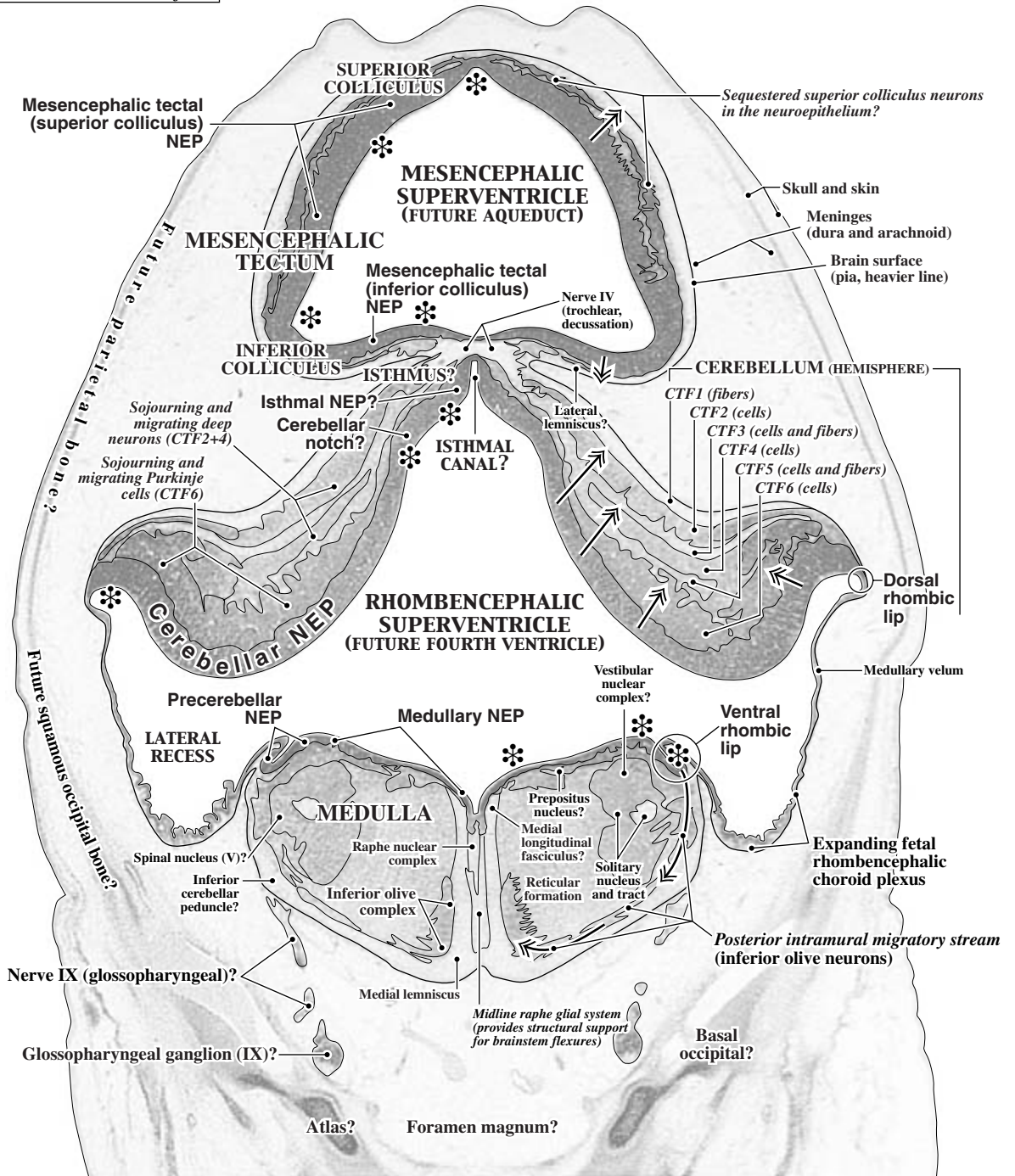
**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 19:
Section 163**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

ABBREVIATIONS:
NEP - Neuroepithelium
CTF - Cerebellar transitional field

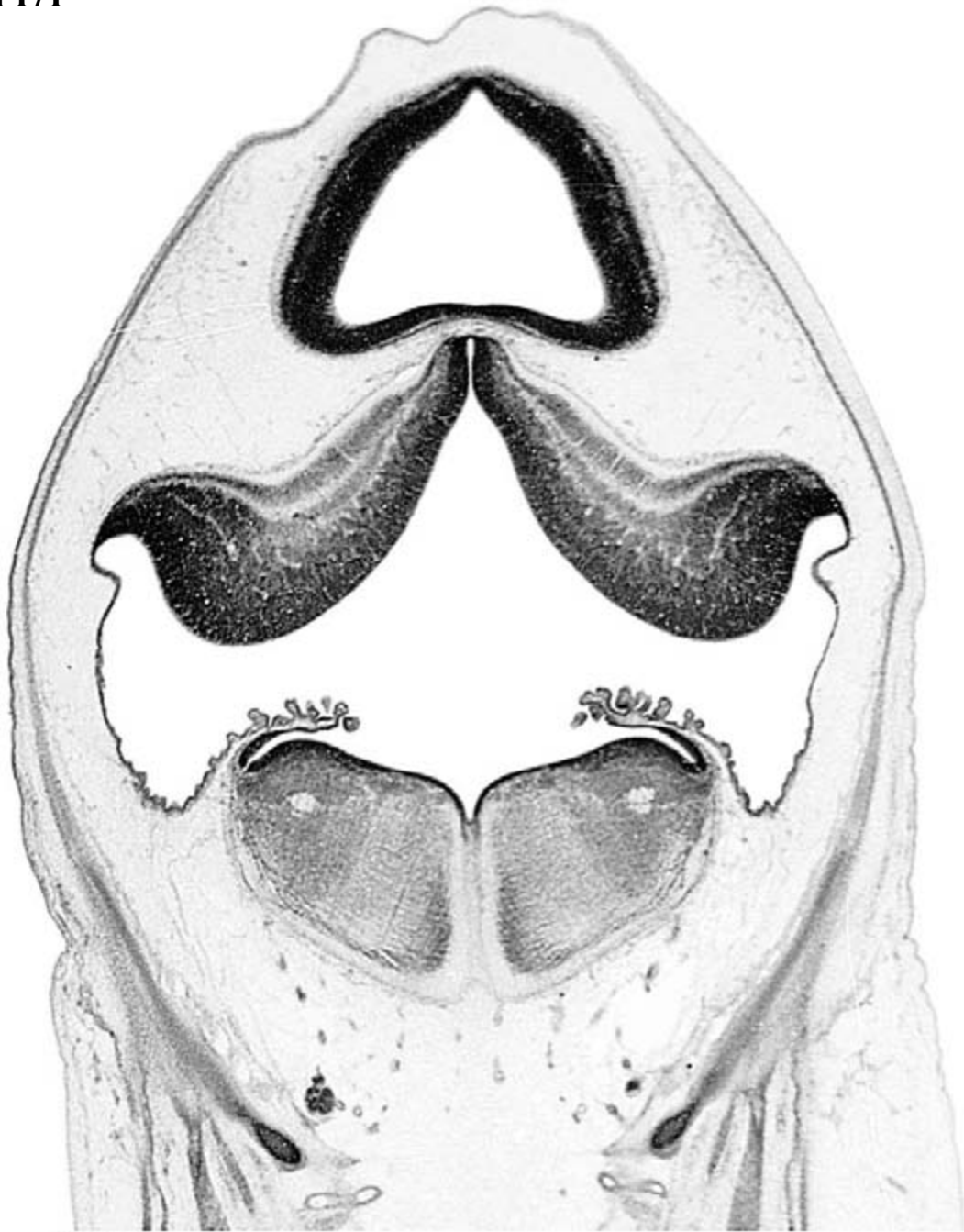


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

PLATE 205A

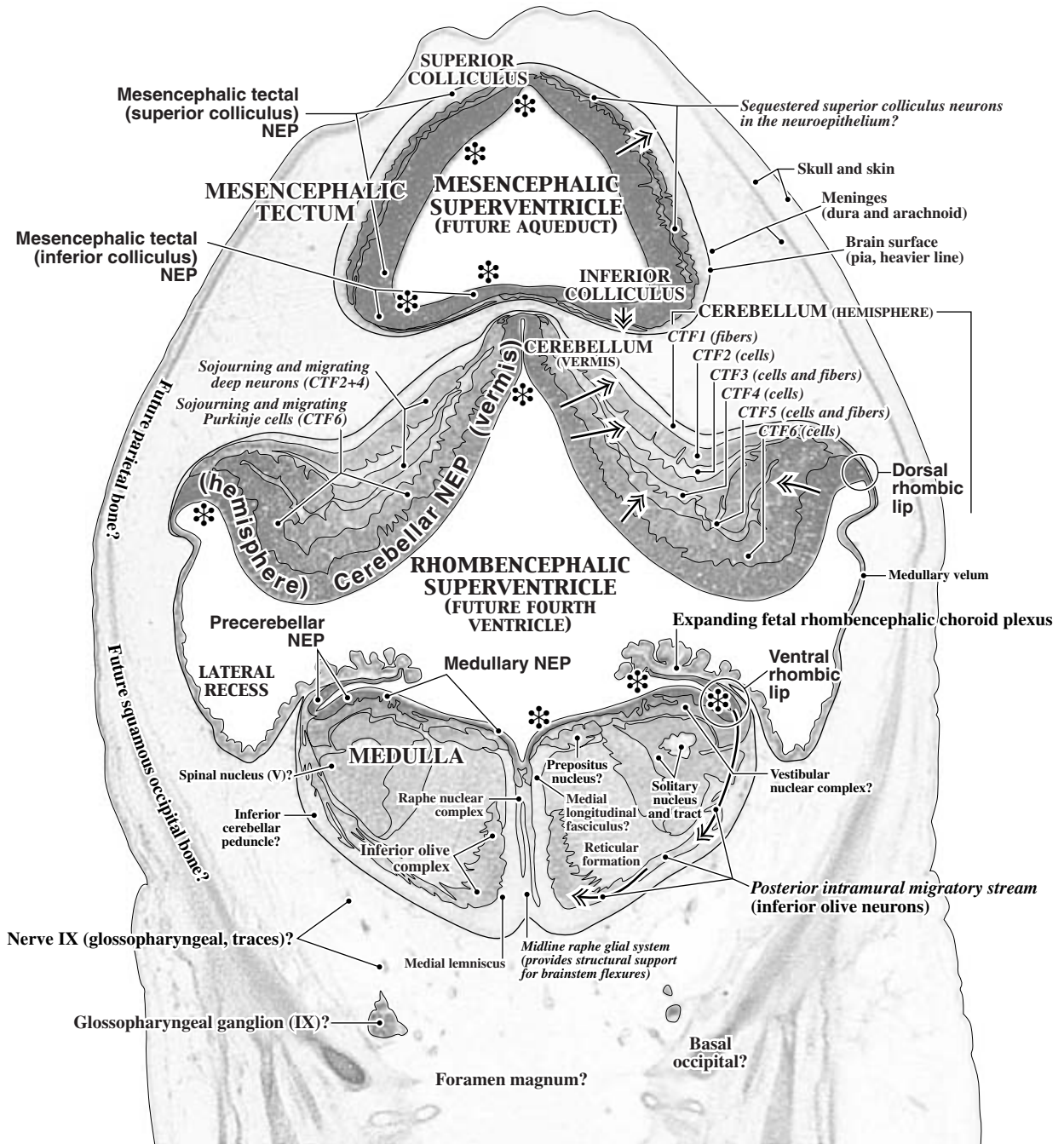
**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 20:
Section 171**



1 mm

FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
NEP - Neuroepithelium
CTF - Cerebellar transitional field



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

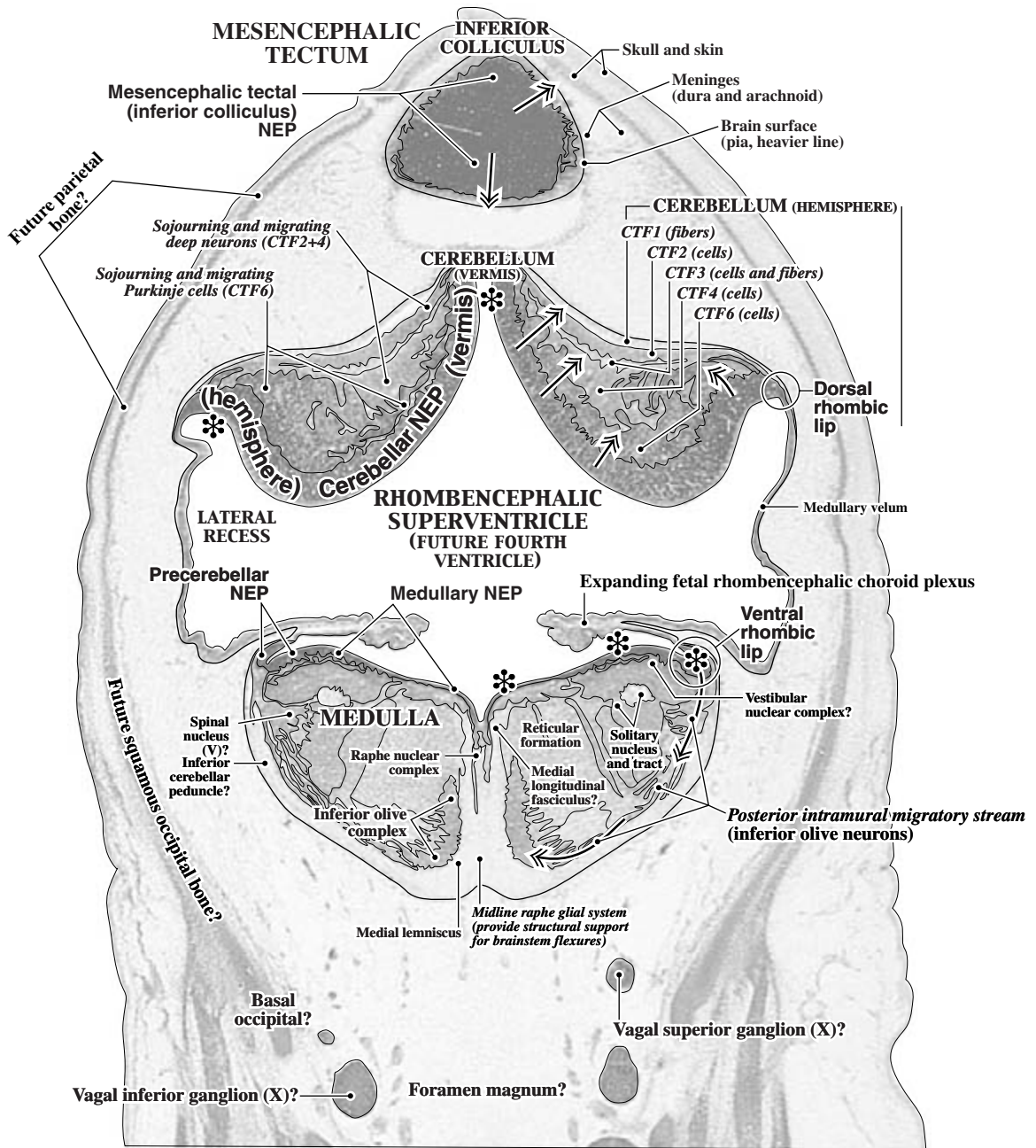
PLATE 206A

**GW7.5 Coronal/horizontal
CR 23 mm, C966
Level 21:
Section 179**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
Germinal zone - Helvetica bold
Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

ABBREVIATIONS:
NEP - Neuroepithelium
CTF - Cerebellar transitional field

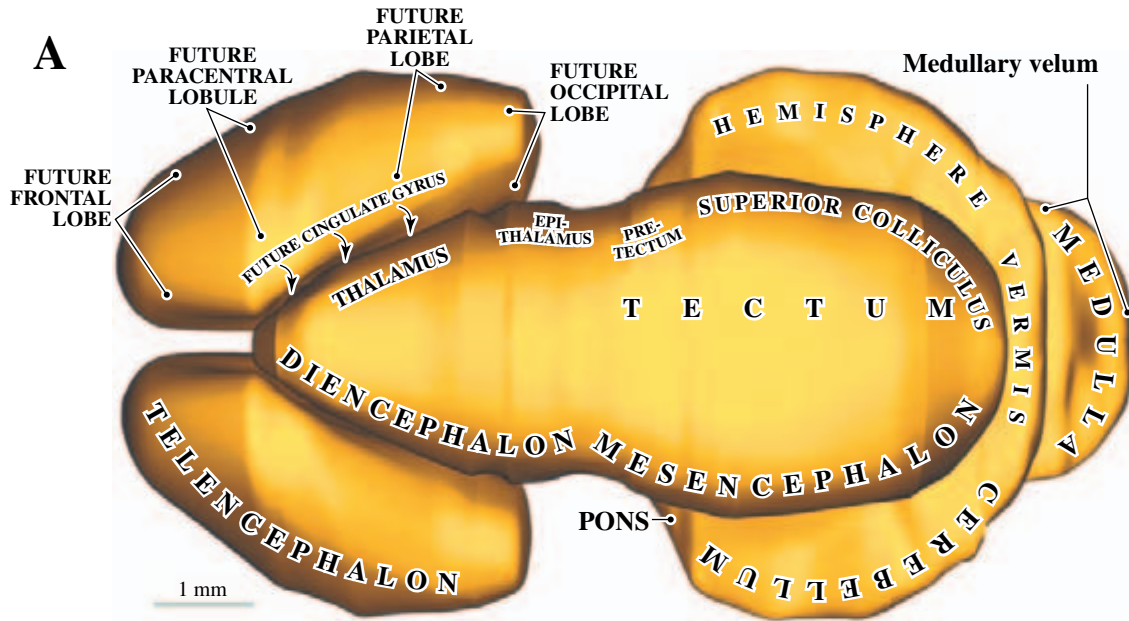


↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

* Evaginations and invaginations of the neuroepithelium are mosaic compartments that give rise to different brain structures.

FIGURE 10

GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION



A. TOP VIEW OF THE BRAIN SURFACE

The observer is looking straight down on the top of the brain. Note the small size of the telencephalon. In a mature brain, the only structure visible from the top is the telencephalon. At GW 7.5, the telencephalon does not yet cover the dorsal surface of the diencephalon. All of the mesencephalic tectum is visible. The cerebellum appears as a wide ledge forming a horseshoe-shaped understory beneath the tectum. The pons is only partly visible connecting to the anteroventral edge of the cerebellum. The medullary velum is all that is visible of the medulla.

B. TOP VIEW SHOWING THE SUPERVENTRICLES

A substantial portion of the brain's volume is occupied by the ventricles. In a mature brain, the ventricles are small, central cavities. At GW 7.5, the diencephalic and mesencephalic superventricles are already narrowing to resemble their adult shapes, and the brain wall is thickest in that region. Most of the telencephalon is filled with the paired telencephalic superventricles, but note that the brain wall is thicker laterally than medially in accordance with the ventrolateral (earlier maturing) to dorsomedial (later maturing) developmental gradient. The dark areas within the brain wall are caused by looking through more than one thickness.

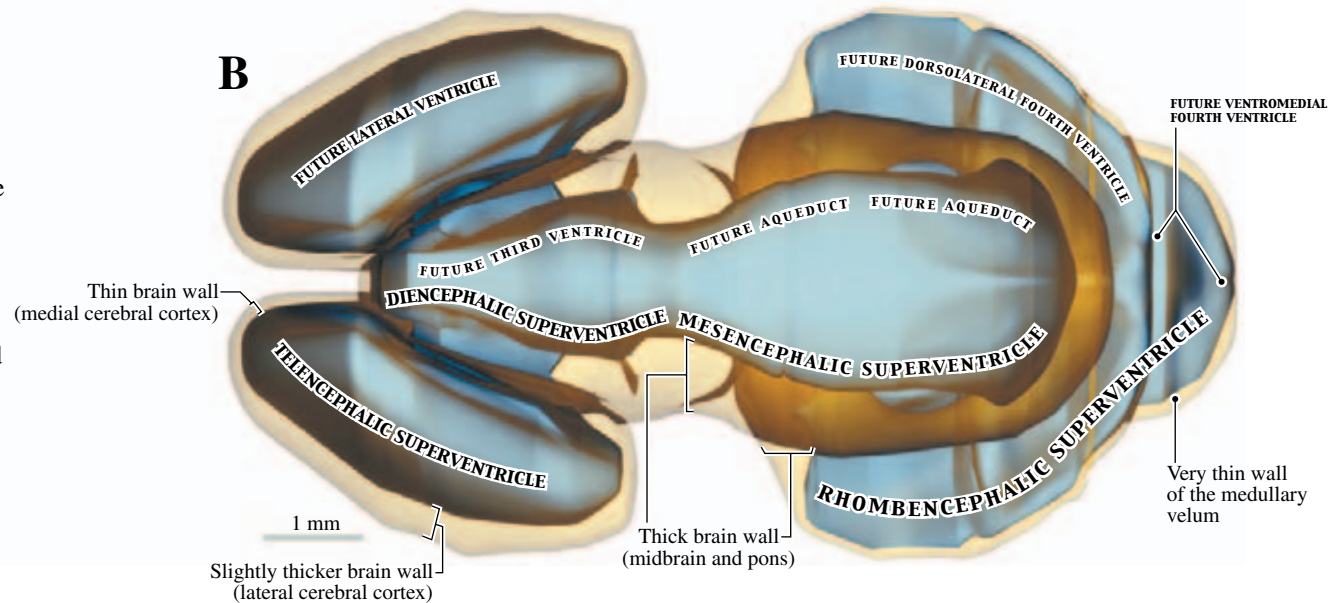
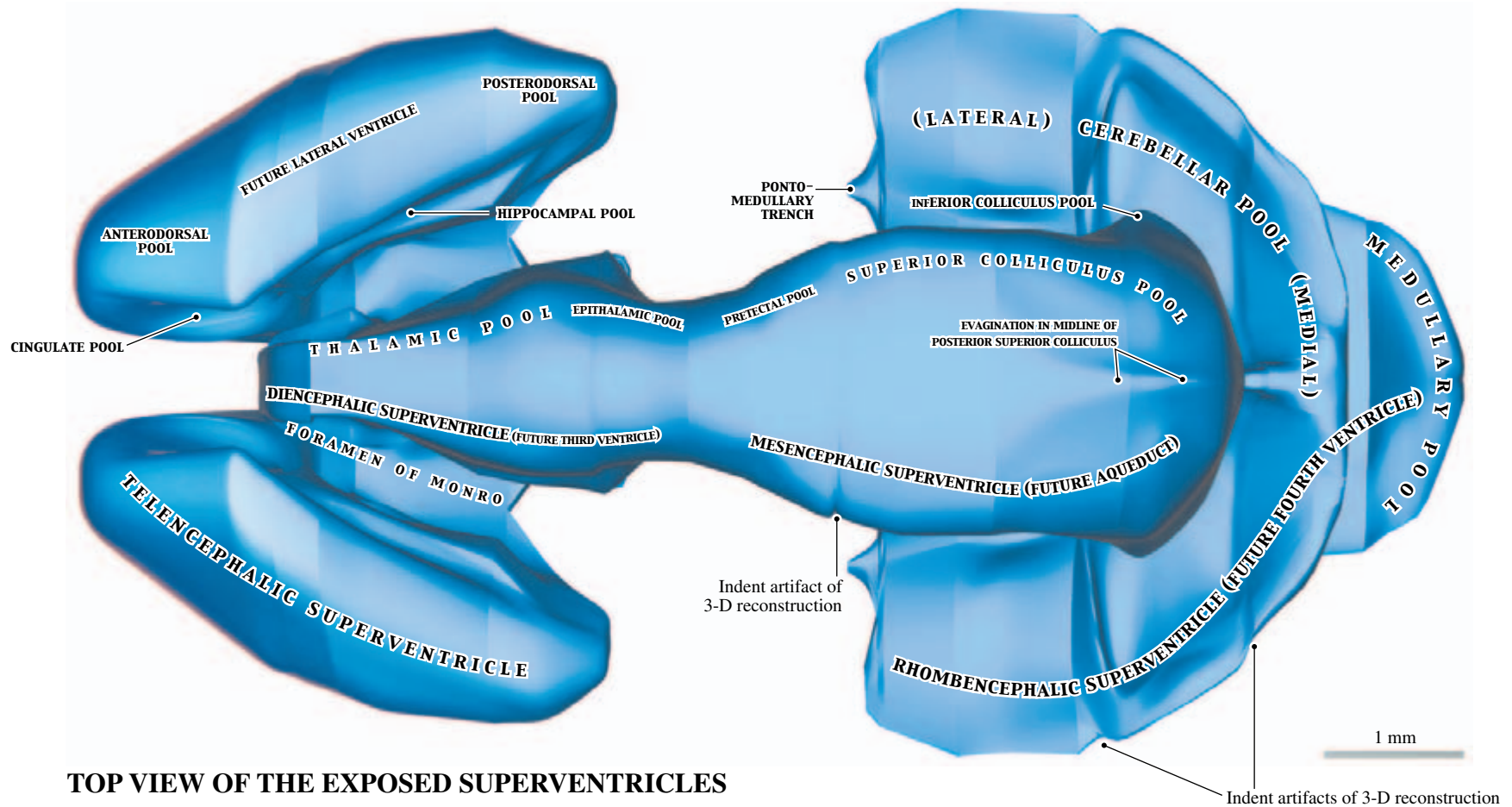


FIGURE 11

GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION

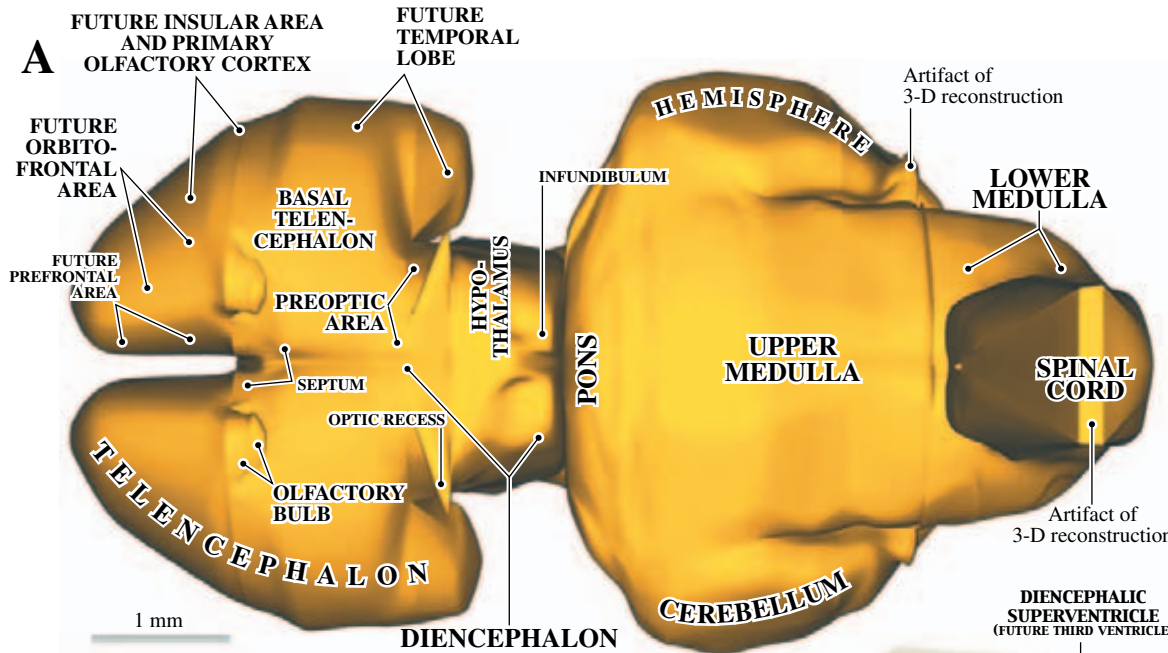


TOP VIEW OF THE EXPOSED SUPERVENTRICLES

Enlarged view of **Figure 10B** showing only the ventricles. Because the ventricles contain fluid, subdivisions are called "pools." The various pools are named according to adjacent structures, which often produce evaginations into the ventricle (such as the cingulate and hippocampal pools). The complex shape of the brain ventricles is primarily a result of (1) localized differential proliferation in the adjacent neuroepithelium and (2) flexures in the diencephalon, midbrain, pons, and medulla during development.

FIGURE 12

GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION



A. BOTTOM VIEW OF THE BRAIN SURFACE

The observer is below, looking straight up at the bottom of the brain and spinal cord. Note the olfactory bulb is located behind the cortex and evaginates from the basal telencephalic area (just above the developing olfactory epithelium in the skull, see **Plates 188 and 189**). The enormous growth of the prefrontal area of the cortex eventually displaces the bulb to lie in a more anterior position. This ventral view shows little of the ventral mesencephalon, because the tegmentum is folded above the ventral diencephalon. The cerebellum still appears as a wide ledge on the lateral edge of the pons that arches over the upper and lower medulla. The spinal cord is coming straight down from the lower medulla.

B. BOTTOM VIEW SHOWING THE SUPERVENTRICLES

The brain wall is thickest in the pons from this viewpoint because the pontine parenchyma is in front of the anterior edge of the rhombencephalic superventricle. In the telencephalon, the lateral brain wall is thicker than the medial wall in accordance with the ventrolateral (earlier maturing) to dorsomedial (later maturing) developmental gradient. As in **Figure 10B**, the dark areas within the brain wall are caused by looking through more than one thickness.

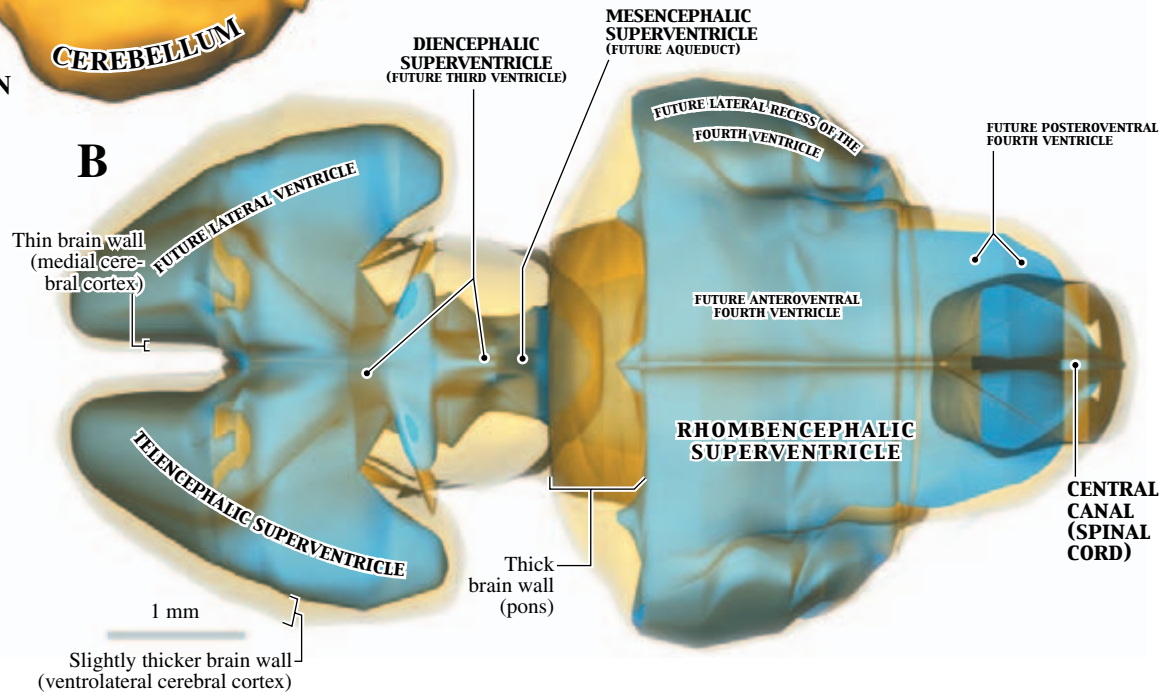
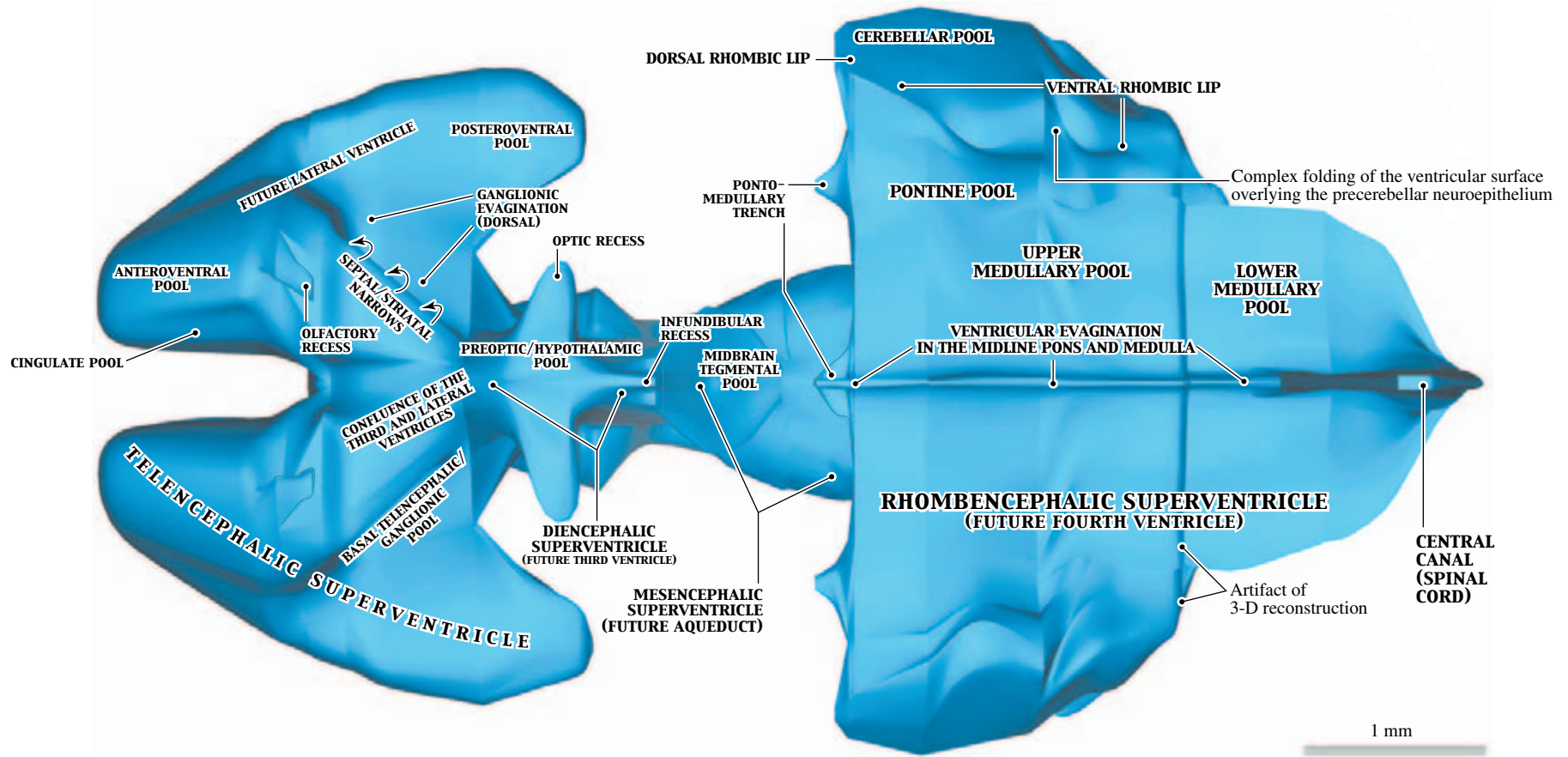


FIGURE 13

GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION



BOTTOM VIEW OF THE EXPOSED SUPERVENTRICLES

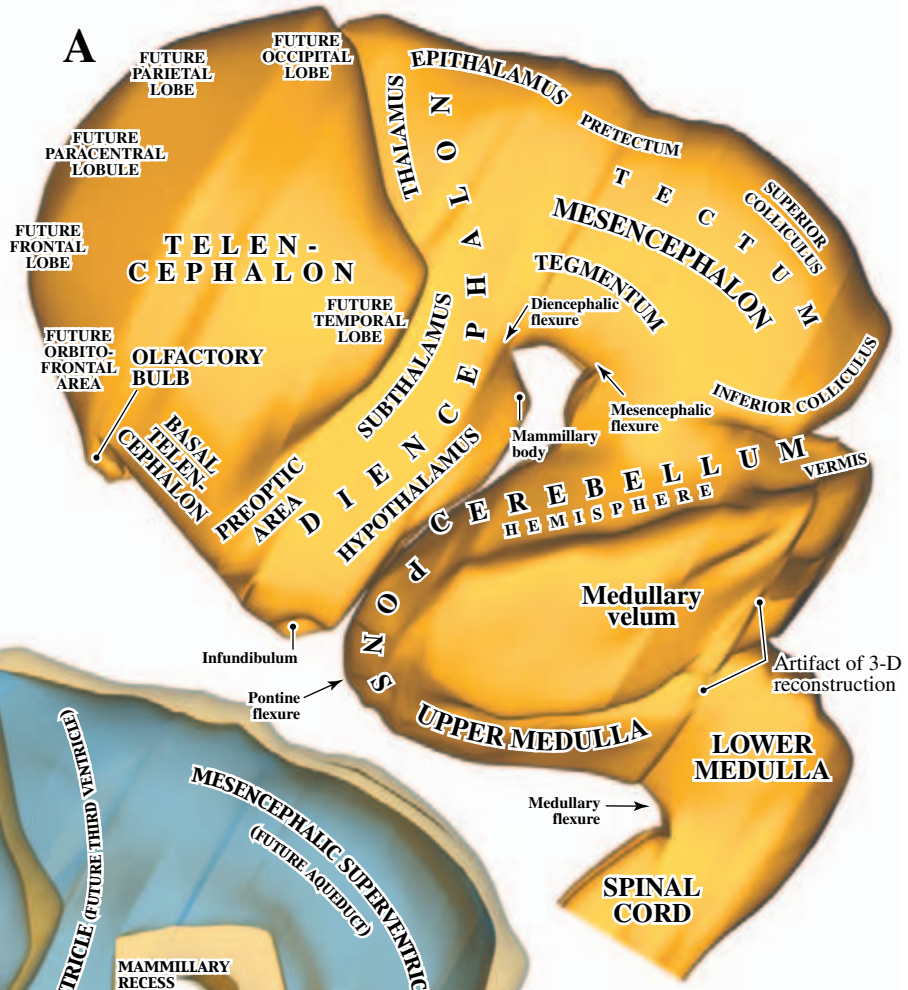
Enlarged view of **Figure 12B** showing only the ventricles. The complex evaginations of the ventricular surface indicate the heterogeneous nature of the neuroepithelium that lines different ventricular divisions.

FIGURE 14

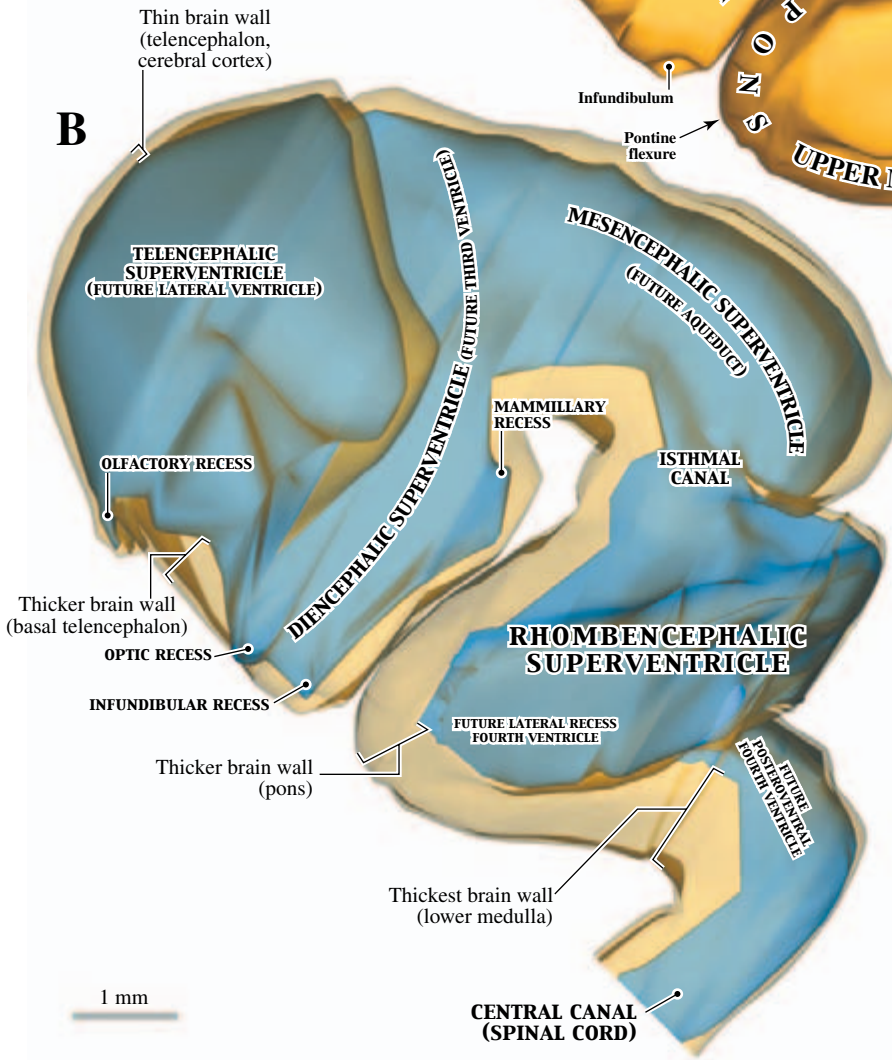
**GW7.5, CR23 mm, C966,
COMPUTER-AIDED
3-D RECONSTRUCTION**

**A. SIDE VIEW OF
THE BRAIN SURFACE**

The observer is beside the model, looking at the lateral surface of the brain and upper cervical spinal cord. Note the posterior-facing olfactory bulb beneath the basal telencephaon. This lateral view shows most clearly all of the flexures in the brainstem.



B



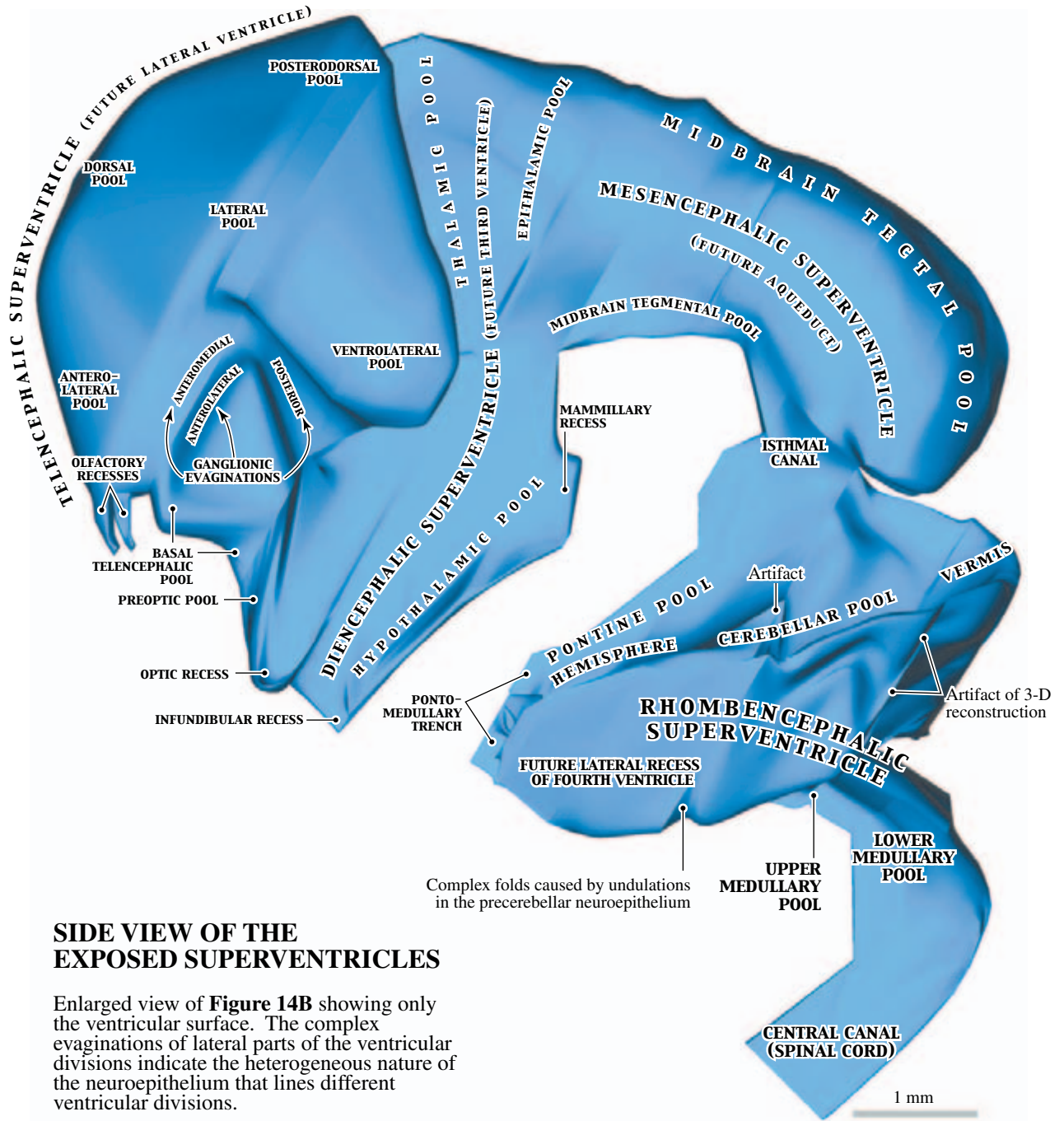
**B. SIDE VIEW
SHOWING THE
SUPERVENTRICLES**

The varying thicknesses of the brain wall mirror a maturation gradient: the medulla has the thickest wall and is most mature, while the thin wall of the cerebral cortex is one of the least mature brain areas. As in **Figures 10B and 12B**, the dark areas within the brain wall are caused by looking through more than one thickness.

1 mm

**GW7.5, CR23 mm, C966,
COMPUTER-AIDED
3-D RECONSTRUCTION**

FIGURE 15

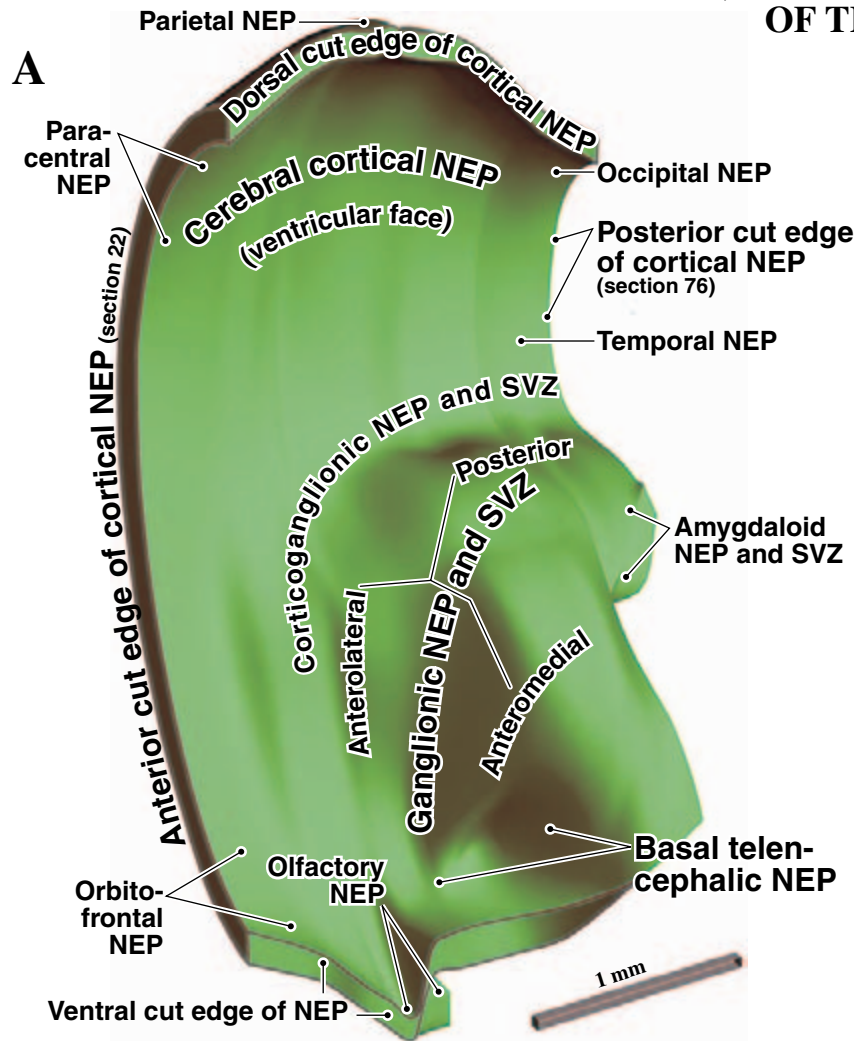


**SIDE VIEW OF THE
EXPOSED SUPERVENTRICLES**

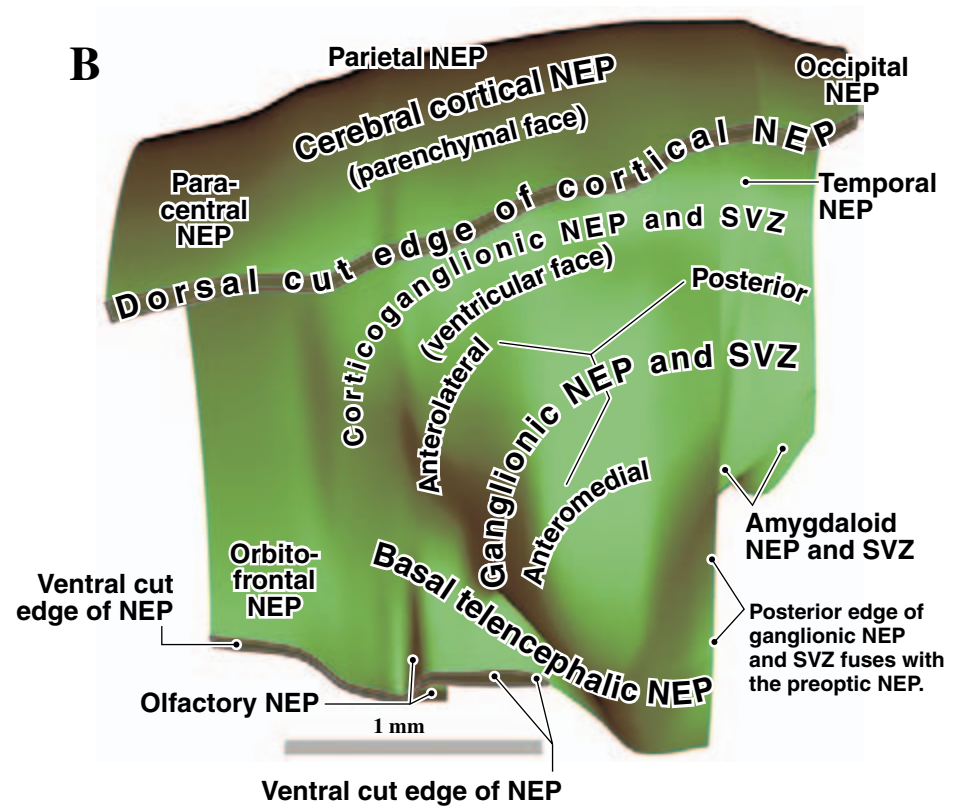
Enlarged view of **Figure 14B** showing only the ventricular surface. The complex evaginations of lateral parts of the ventricular divisions indicate the heterogeneous nature of the neuroepithelium that lines different ventricular divisions.

FIGURE 16

GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION OF THE LATERAL TELEENCEPHALIC NEUROEPITHELIUM



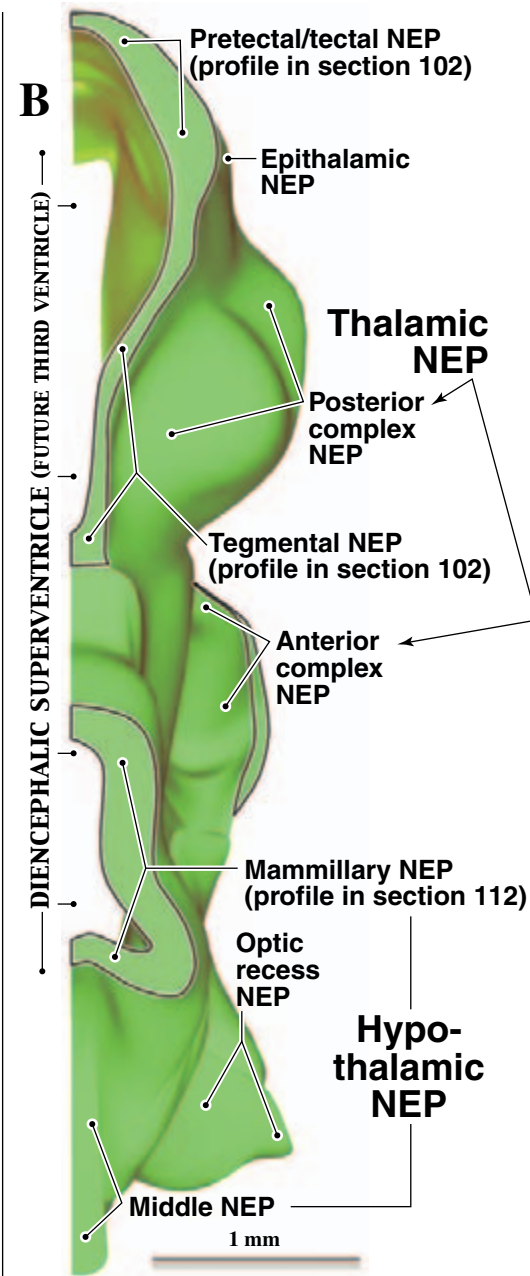
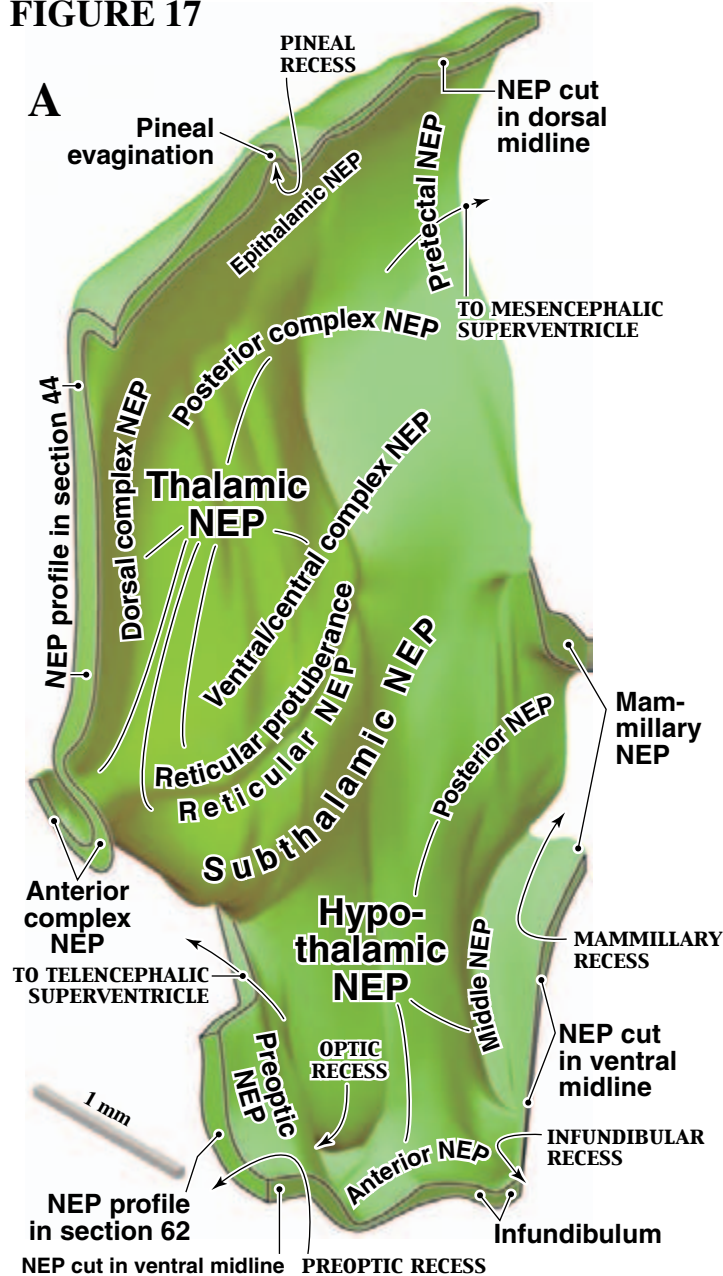
A. is an inside "camera" view of the lateral half of the telencephalic neuroepithelium and subventricular zone; the medial half is not reconstructed so that the ganglionic and olfactory evaginations are visible. The reconstruction includes sections 22 (between Levels 1 and 2, Plates 187 and 188) through 76 (Level 7, Plate 192). The camera is looking straight (0° pitch) at the reconstruction with a +60° rotation (heading). The large ganglionic evagination is evident in the ventrolateral NEP and SVZ.



B. is another inside "camera" view of the same reconstruction in A. The camera is looking down (-70° pitch) at the reconstruction with a +90° rotation (heading). From that angle, the superficial face of the dorsal cortical NEP is visible, and the large ganglionic evagination rises up into the future lateral ventricle. The posterior edge of the ganglionic NEP and SVZ fuses with the preoptic NEP posterior to the foramen of Monro. See that transition between Level 5 (Plate 190) and Level 6 (Plate 191).

<p>ABBREVIATIONS: NEP - Neuroepithelium SVZ - Subventricular zone</p>	<p>Neuroepithelial compartments labeled in Helvetica Bold.</p>
--	---

FIGURE 17



**GW7.5, CR23 mm, C966,
COMPUTER-AIDED
3-D RECONSTRUCTION
OF THE DIENCEPHALIC
NEUROEPITHELIUM**

Two inside "camera" views of the right half of the diencephalic neuroepithelium that includes section 44 (near **Level 3, Plate 188**) through section 112 (**Level 12, Plate 197**); the NEP in all sections has been cut where it bridges the midline dorsally and ventrally so that its folds and undulations can be observed.

A. The camera views the front of the reconstruction with +45° heading and -25° pitch. From this angle, we see the ventricular face of the NEP.

B. The camera views the back of the reconstruction with 180° heading and -10° pitch. From this angle, we see some of the parenchymal face of the NEP. Note the prominent folding in the anterior complex NEP, the thick posterior complex NEP, and the back wall of the NEP lining the optic recess.

It is postulated that the multiple evaginations and invaginations of the diencephalic neuroepithelium are mosaic compartments that give rise to different thalamic, hypothalamic, and preoptic nuclei.

ABBREVIATION:
NEP - Neuroepithelium

Neuroepithelial compartments
labeled in Helvetica Bold.
VENTRICULAR DIVISIONS
LABELED IN CAPITALS.

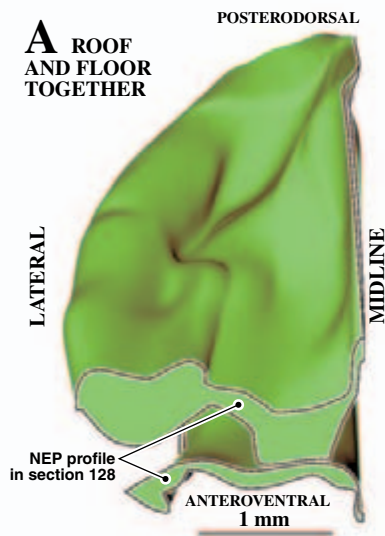
FIGURE 18
GW7.5, CR23 mm, C966,
COMPUTER-AIDED
3-D RECONSTRUCTION OF
THE RHOMBENCEPHALIC
NEUROEPITHELIUM

The reconstruction includes section 128 (Level 15, Plate 200) through section 193 (posterior to Level 21, Plate 179); the NEP in all sections has been cut where it bridges the midline dorsally and ventrally, and only the right half is shown. The camera is looking at the front of the NEP (0° heading, -10° pitch).

A. shows both the roof and floor plates of the rhombencephalic NEP with orientation labels.

B. shows the parenchymal face of the roof NEP bordering the differentiating zones of the pons, isthmus, and cerebellum. Since the edges between *CTF6* (Purkinje cell sojourn zone) and the **cerebellar NEP** are virtually indistinguishable, both are included in the reconstruction.

C. shows the ventricular face of the floor NEP overlying the upper (adjacent to pons) and lower medulla (adjacent to spinal cord).

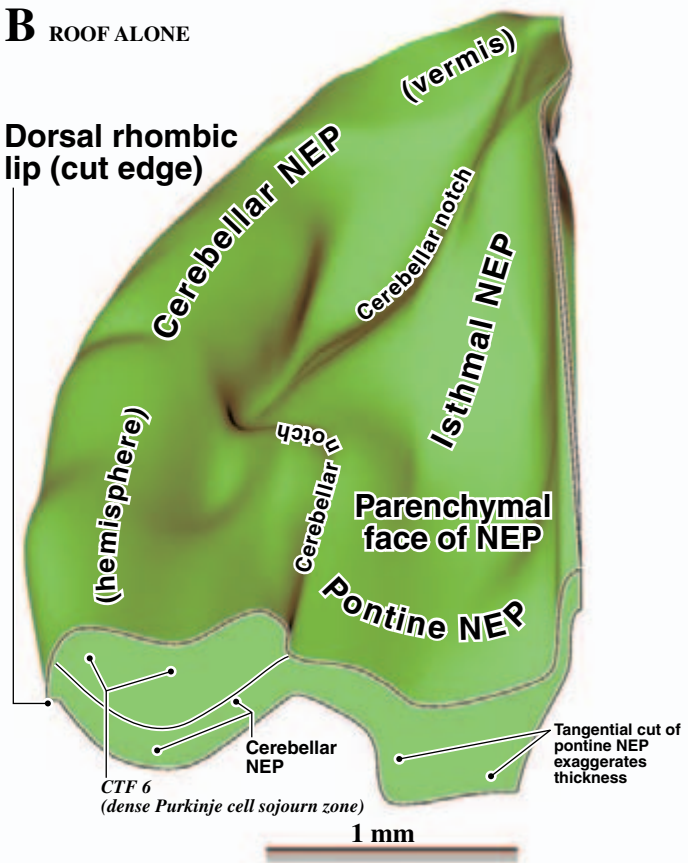


ABBREVIATIONS:
CTF - Cerebellar transitional field
 NEP - Neuroepithelium

Neuroepithelial compartments
labeled in Helvetica Bold.
Transient developmental structures
labeled in Times Bold Italic

B ROOF ALONE

Dorsal rhombic lip (cut edge)



C FLOOR ALONE

Midline medullary NEP evagination

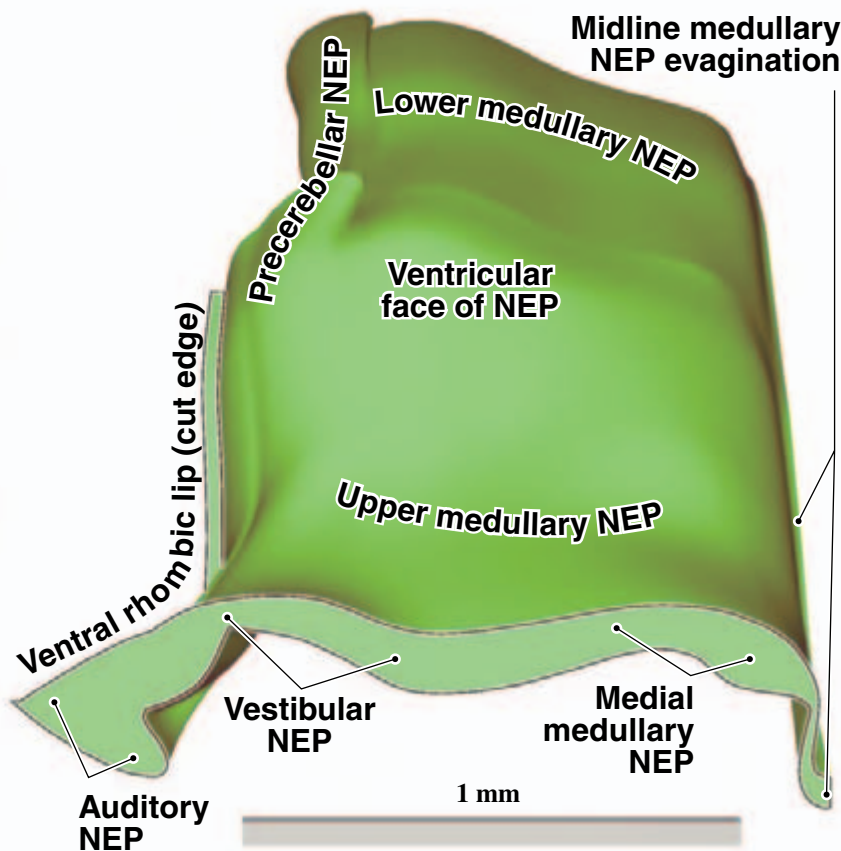
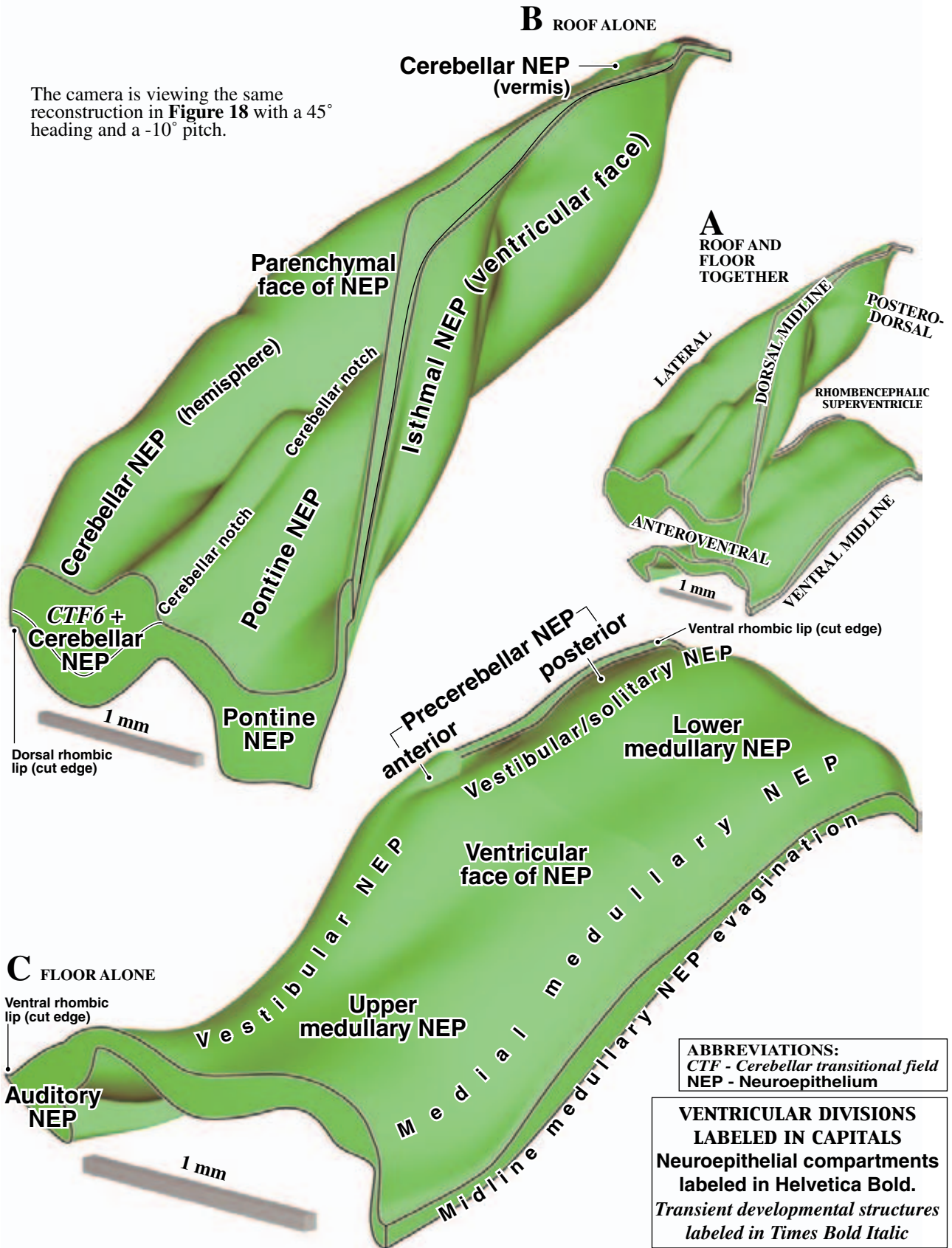


FIGURE 19
GW7.5, CR23 mm, C966, COMPUTER-AIDED 3-D RECONSTRUCTION
OF THE RHOMBENCEPHALIC NEUROEPITHELIUM



PART XI: GW7.5 SAGITTAL

This specimen number 6202 in the Carnegie Collection, designated here as C6202, is a fetus of unknown sex with a crown-rump length (CR) of 21 mm estimated to be at gestational week (GW) 7.5. The entire fetus was cut in the sagittal plane in 20 μ m sections and stained with hematoxylin and eosin. Information on the date of specimen collection, fixative, and embedding medium was not available to us. Since there is no photograph of this specimen before histological processing, a specimen from Hochstetter (1919) that is comparable in age to C6202 is used to show external brain features at GW7.5 (A, **Figure 20**). Like all other sagittal specimens in this Volume, C6202's sections are not cut parallel to the midline; **Figure 20** shows the approximate rotations in horizontal (B) and vertical (C) dimensions. Photographs of 7 sections (**Levels 1-7**) are illustrated at low magnification in four parts (**Plates 207A-D** through **213A-D**). The A/B parts show the brain in place in the skull; the C/D parts show only the brain (and some peripheral ganglia) at slightly higher magnification. **Plates 214-221** show high-magnification views of various parts of the brain at different levels from the cerebral cortex (**Plate 214**) to the midline raphe glial structure in the midbrain and cervical spinal cord (**Plate 221**). Most of the high-magnification plates are rotated 90° (landscape orientation) to more efficiently use page space.

C6202 is less mature than C966 even though it is only 2 mm shorter in CR length. The *supraventricles* are large in the centers of all brain structures, especially in the telencephalon and rhombencephalon, but the telencephalon is smaller than it is in C966. Sections near the midline show the enormous size of the diencephalic and mesencephalic supraventricles. The respective thicknesses of the *neuroepithelium* (NEP) and parenchyma are keys to determining the degree of maturation of various brain structures.

The parenchyma is thick and bordered by a thin NEP in the medial medulla, indicating that many neurons have been generated here, but the production of late-generated neurons continues. The parenchyma is thinner and the NEP is thicker in the lateral medulla, entire pons, and entire midbrain tegmentum. There are layers of dense cells adjacent to the NEP where neurons are sojourning and the rest of the pontine and mesencephalic tegmental parenchyma is filled with migrating and settling neurons. The *pre-cerebellar neuroepithelium* in the lateral medulla is thick and many neurons are entering the *posterior intramural migratory stream*, but few have accumulated in the indistinct inferior olive in the medial medulla. The cerebellar parenchyma has only four layers in the *cerebellar transitional field (CTF)*. The cells in *CTF2, 3, and 4* are probably deep neurons that will eventually settle in the dentate, interpositus, and fastigial nuclei. *CTF5 and 6* are present in C966 and presumably contain Purkinje cells, but these layers are absent in C6202. The cerebellar NEP is thicker than it is in C966 and is now producing the oldest Purkinje cells. The mesencephalic tectal NEP is very thick and lies adjacent to a thin parenchyma. The majority of neurons in both the superior and inferior colliculi have not yet been generated. The diencephalic NEP is thick and the adjacent parenchyma is filled with dense zones of sojourning and migrating neurons. The basal telencephalic NEP and the basal ganglionic NEP are thick, and the oldest neurons are settling in the adjacent parenchyma. Many neurons in the basal telencephalon have yet to be generated. In the cerebral cortex, the NEP is bordered by a thin primordial plexiform layer that contains the oldest cortical neurons (Cajal-Retzius cells) and subplate neurons. The cortical NEP is expanding and increasing its number of neural stem cells as the telencephalic supraventricle grows; nearly all cortical neurons in layers II-VI have still to be generated.

GW7.5 SAGITTAL

A perfect sagittal cut through the brain bisects the cerebral cortex into two separate hemispheres by passing through the interhemispheric fissure, and does the same in the brainstem by passing through the midline of the ventricles.

Sections of C6202's brain rotate 10.5° counterclockwise from the horizontal midline running through the cerebral cortex and midbrain tectum (top view). C6202's sections are quite close to the vertical midline, rotating only 2.8° counterclockwise (back view). In the sections illustrated on the following pages, the telencephalon and diencephalon (top) are tilted away from the observer, while the medulla and upper spinal cord (bottom) are tilted toward the observer.

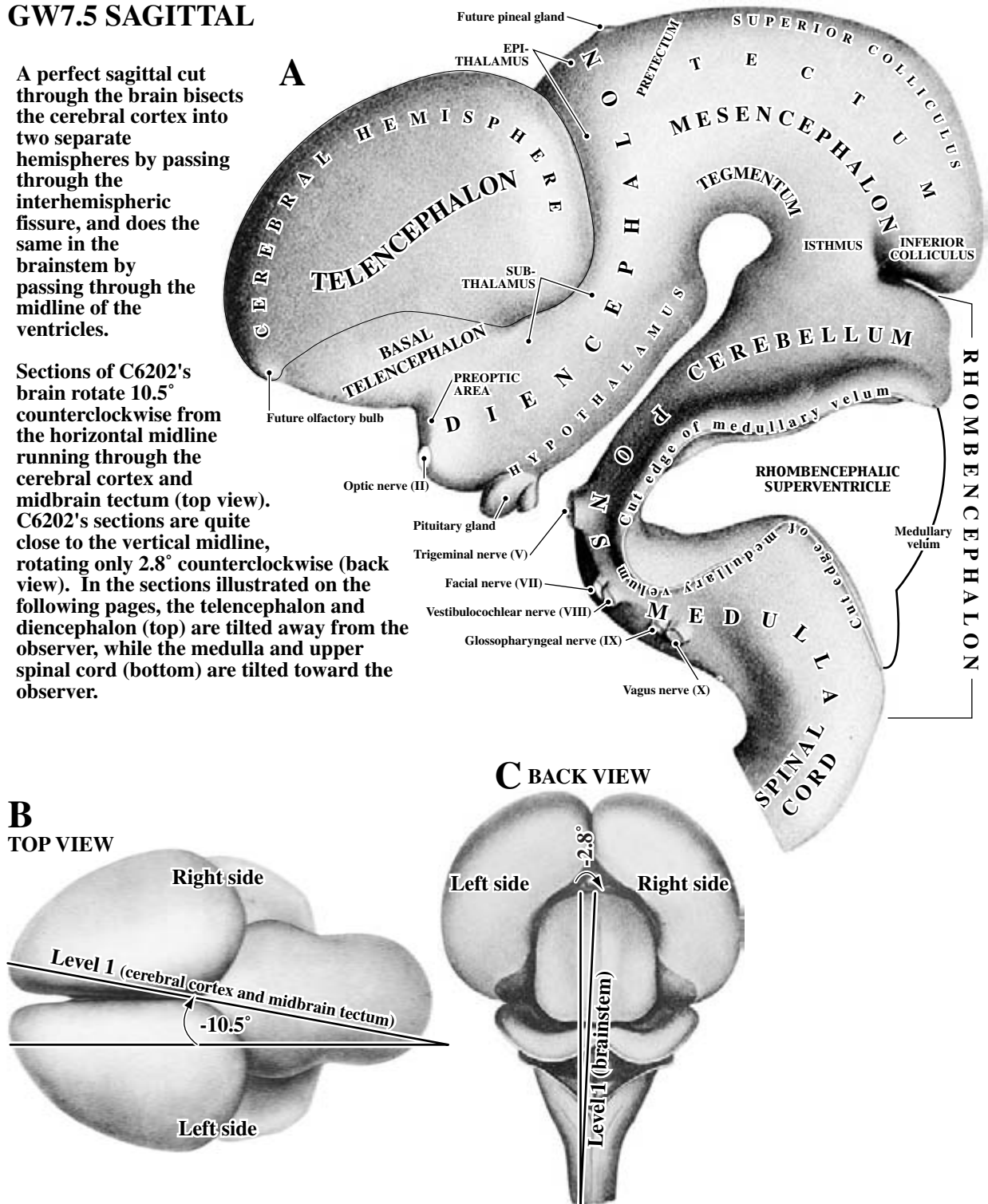
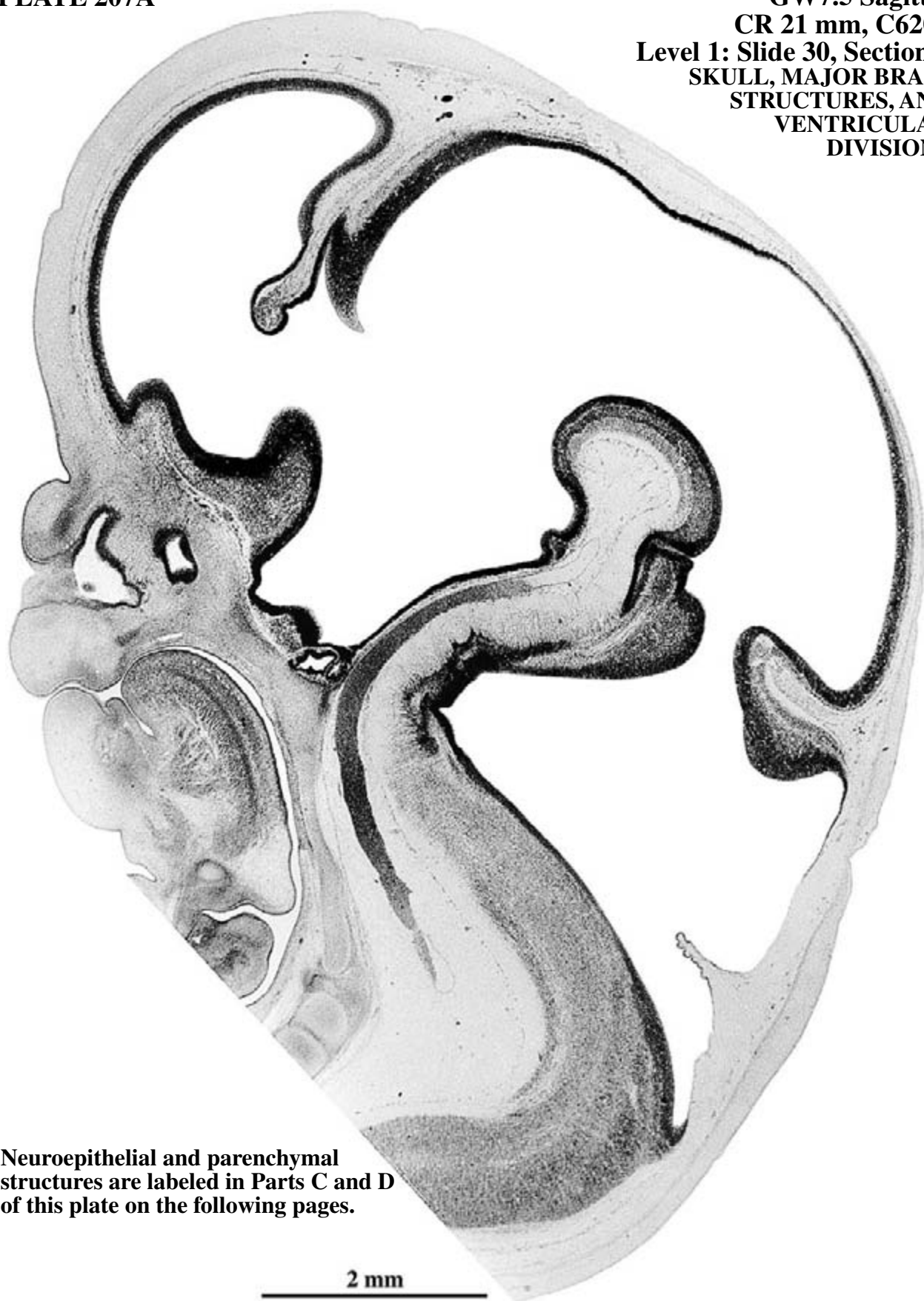


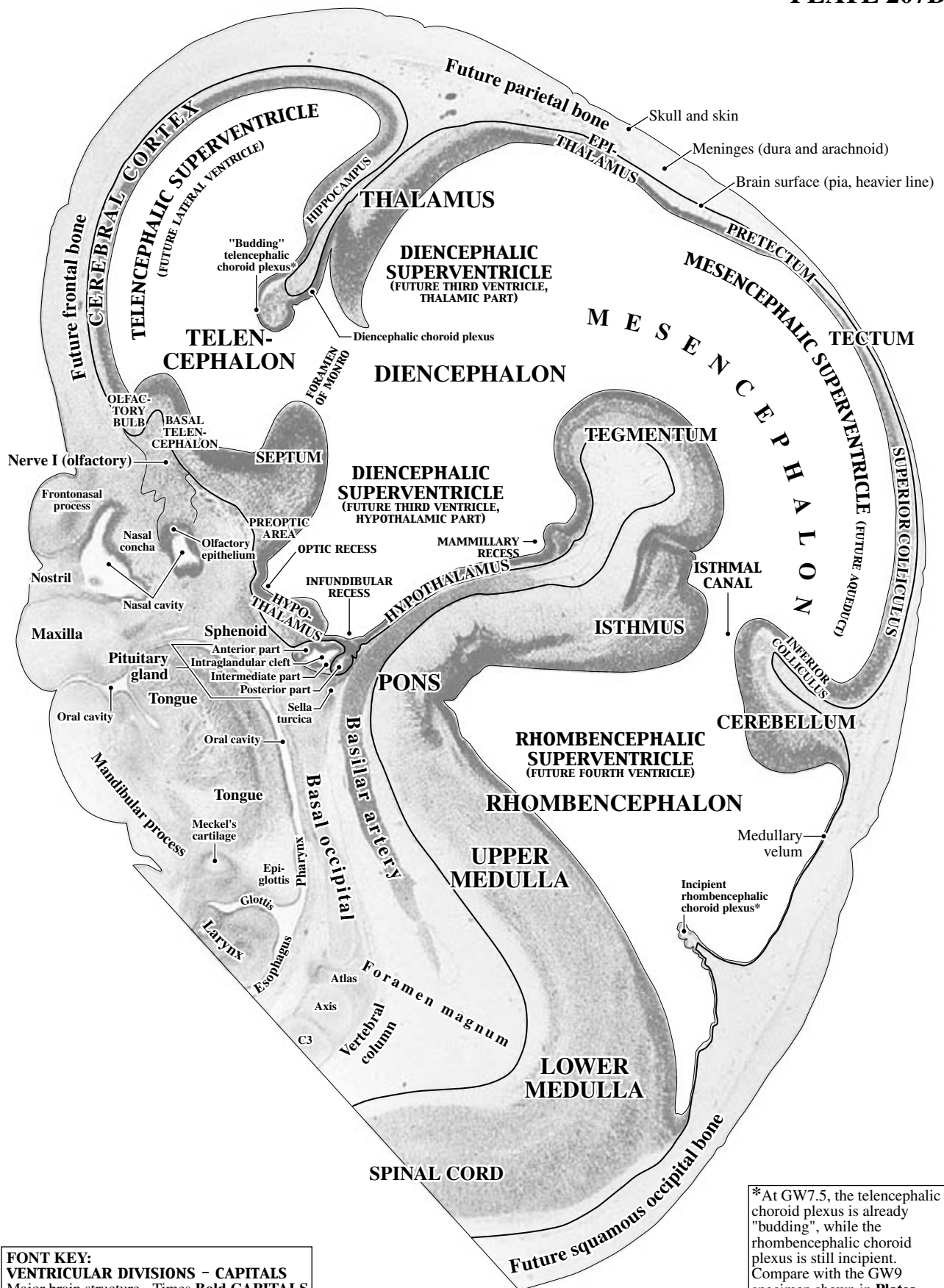
Figure 20. A, The lateral view of the brain and upper cervical spinal cord from a specimen with a crown-rump length of 19.4 mm (modified from Figure 35, Table VI, Hochstetter, 1919) identifies external features of a brain similar to C6202 (CR 21 mm). B, Top view of the brain in A (modified from Figure 37, Table VI, Hochstetter, 1919) shows how C6202's sections rotate from a line parallel to the horizontal midline in the interhemispheric fissure and midbrain tectum. C, Back view of the brain with a crown-rump length of 38 mm (modified from Figure 44, Table VIII, Hochstetter, 1919) shows how C6202's sections rotate from a line parallel to the vertical midline in the brainstem and upper cervical spinal cord.

PLATE 207A

**GW7.5 Sagittal
CR 21 mm, C6202
Level 1: Slide 30, Section 2
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



**Neuroepithelial and parenchymal
structures are labeled in Parts C and D
of this plate on the following pages.**

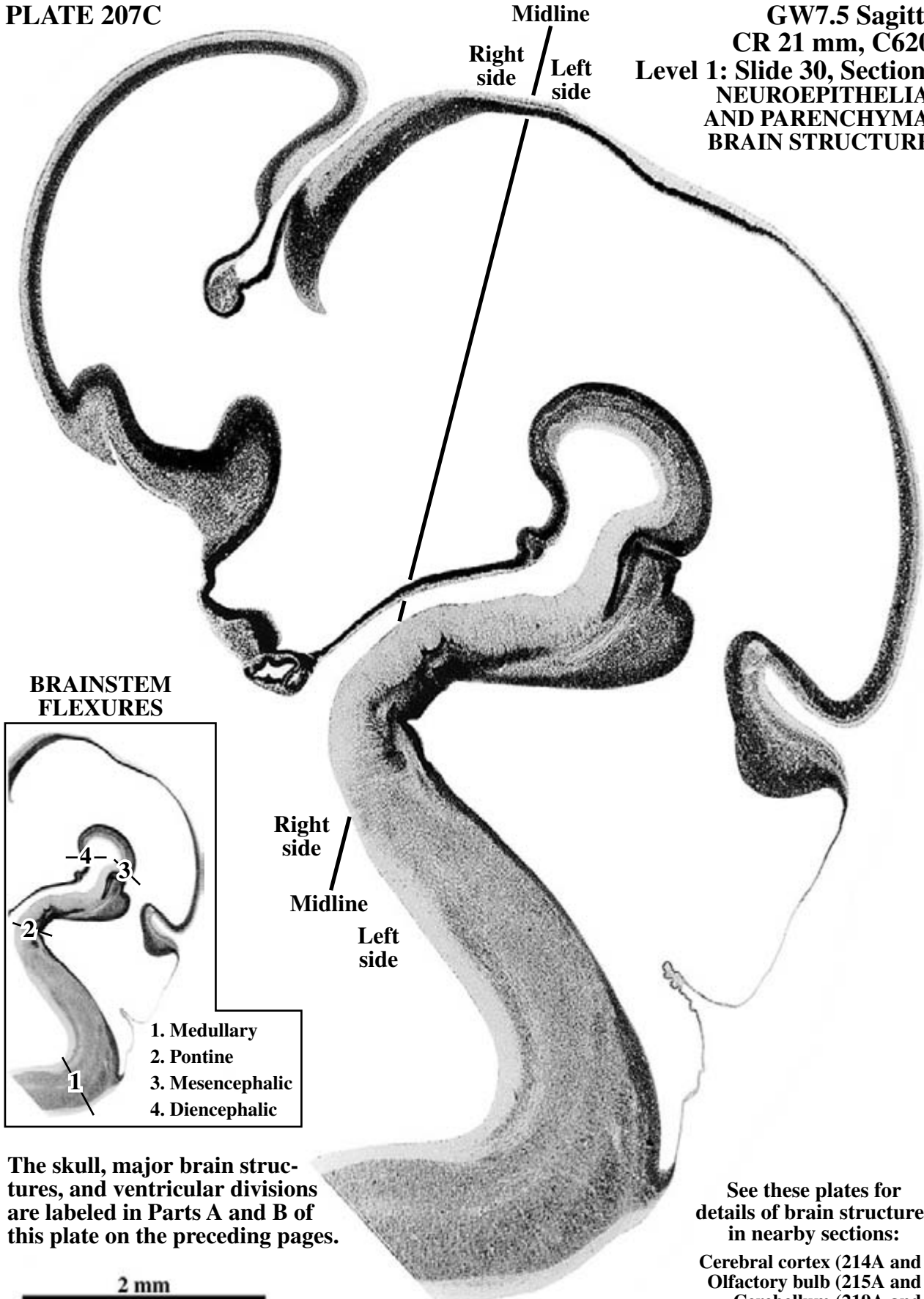


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

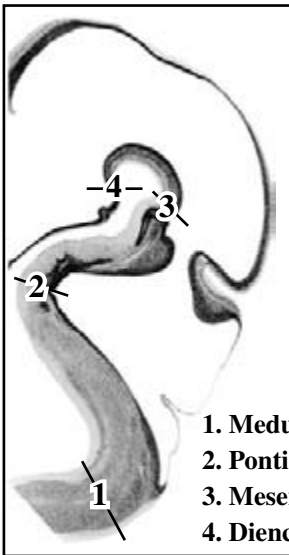
*At GW7.5, the telencephalic choroid plexus is already "budding", while the rhombencephalic choroid plexus is still incipient. Compare with the GW9 specimen shown in Plates 58A and B on pages 137-138.

PLATE 207C

GW7.5 Sagittal
CR 21 mm, C6202
Level 1: Slide 30, Section 2
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES



BRAINSTEM FLEXURES



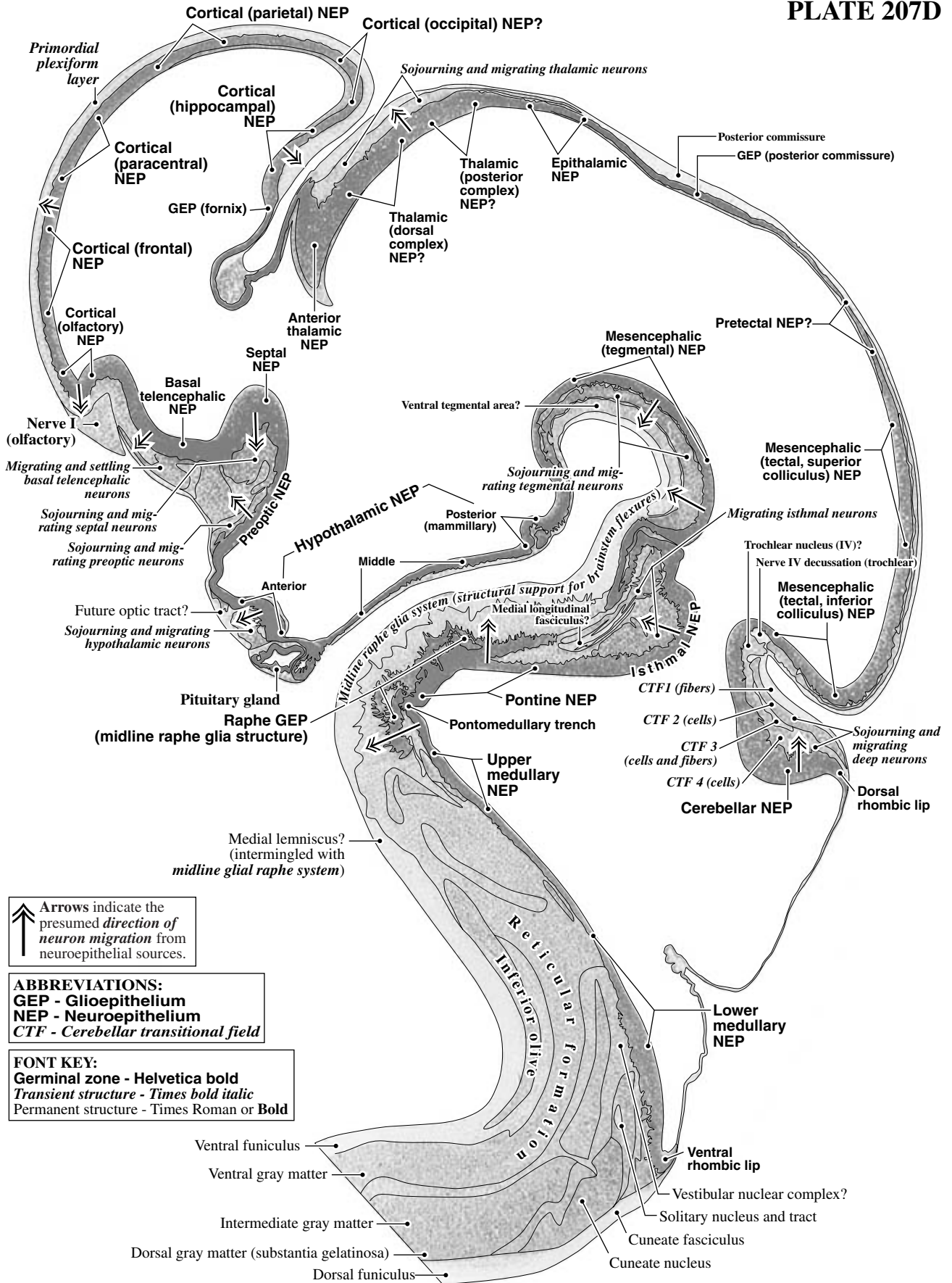
- 1. Medullary
- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

The skull, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

2 mm

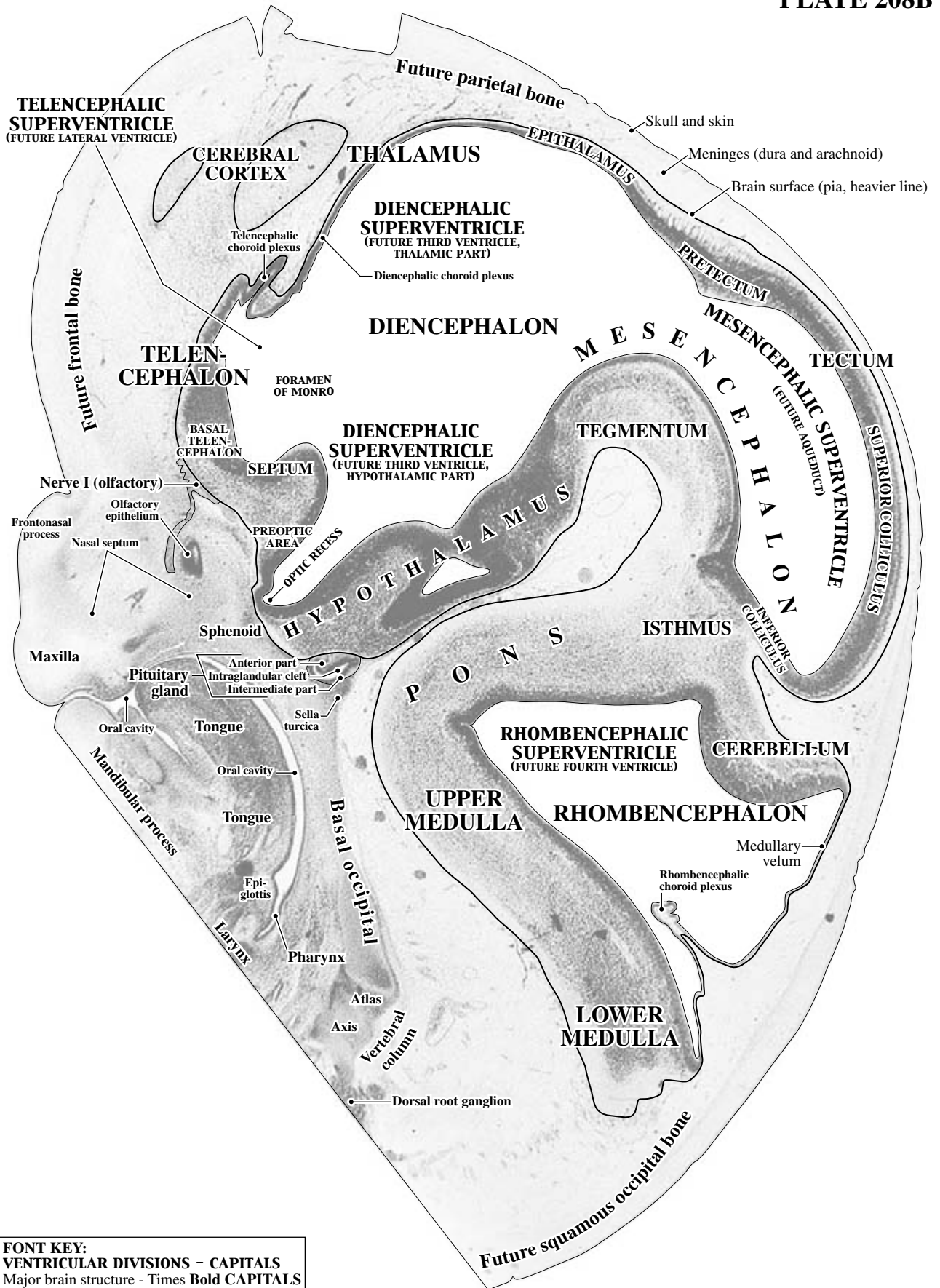
See these plates for details of brain structures in nearby sections:

- Cerebral cortex (214A and B)
- Olfactory bulb (215A and B)
- Cerebellum (219A and B)
- Midline glial fibers (221A and B)



**GW7.5 Sagittal
CR 21 mm, C6202
Level 2: Slide 26, Section 2
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**

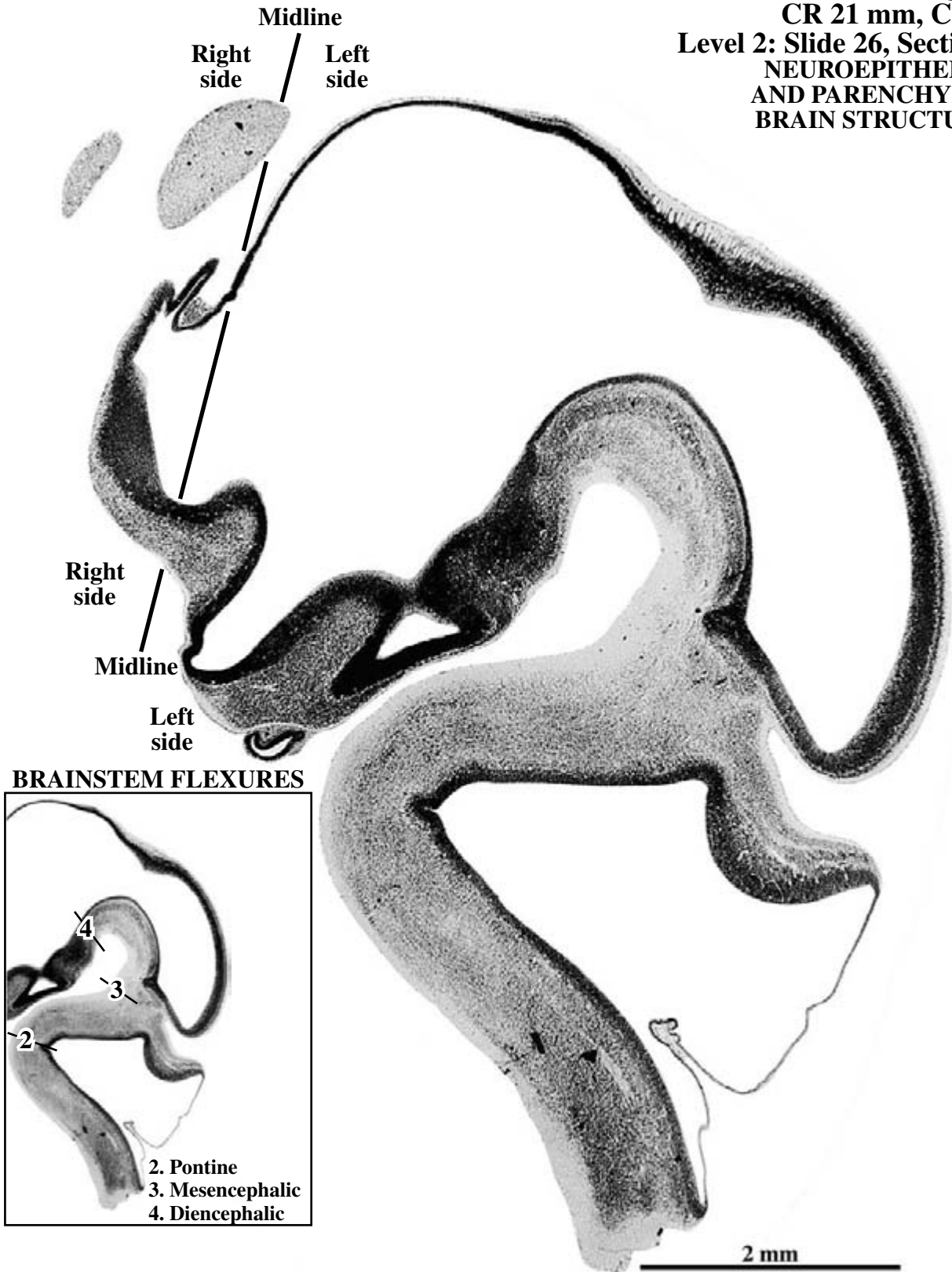




FONT KEY:
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 All other structures - Times Roman or **Bold**

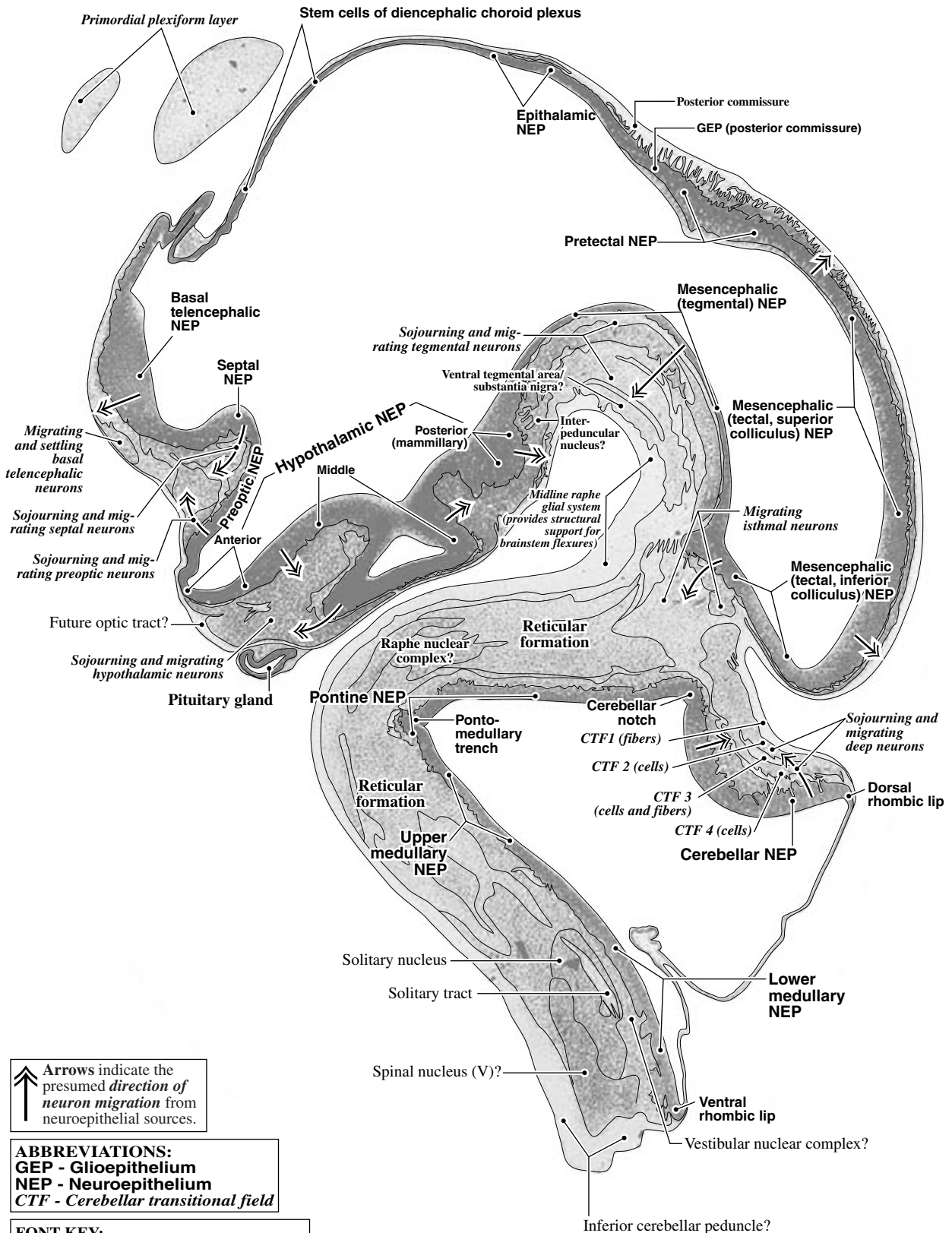
PLATE 208C

GW7.5 Sagittal
CR 21 mm, C6202
Level 2: Slide 26, Section 2
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES



The skull, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.

See Plates 210A and B for details of the anterior and middle hypothalamus, Plates 218A and B for details of the midbrain tegmentum in nearby sections.



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
CTF - Cerebellar transitional field

FONT KEY:
Germinal zone - Helvetica bold
Transient structure - Times bold italic
 Permanent structure - Times Roman or Bold

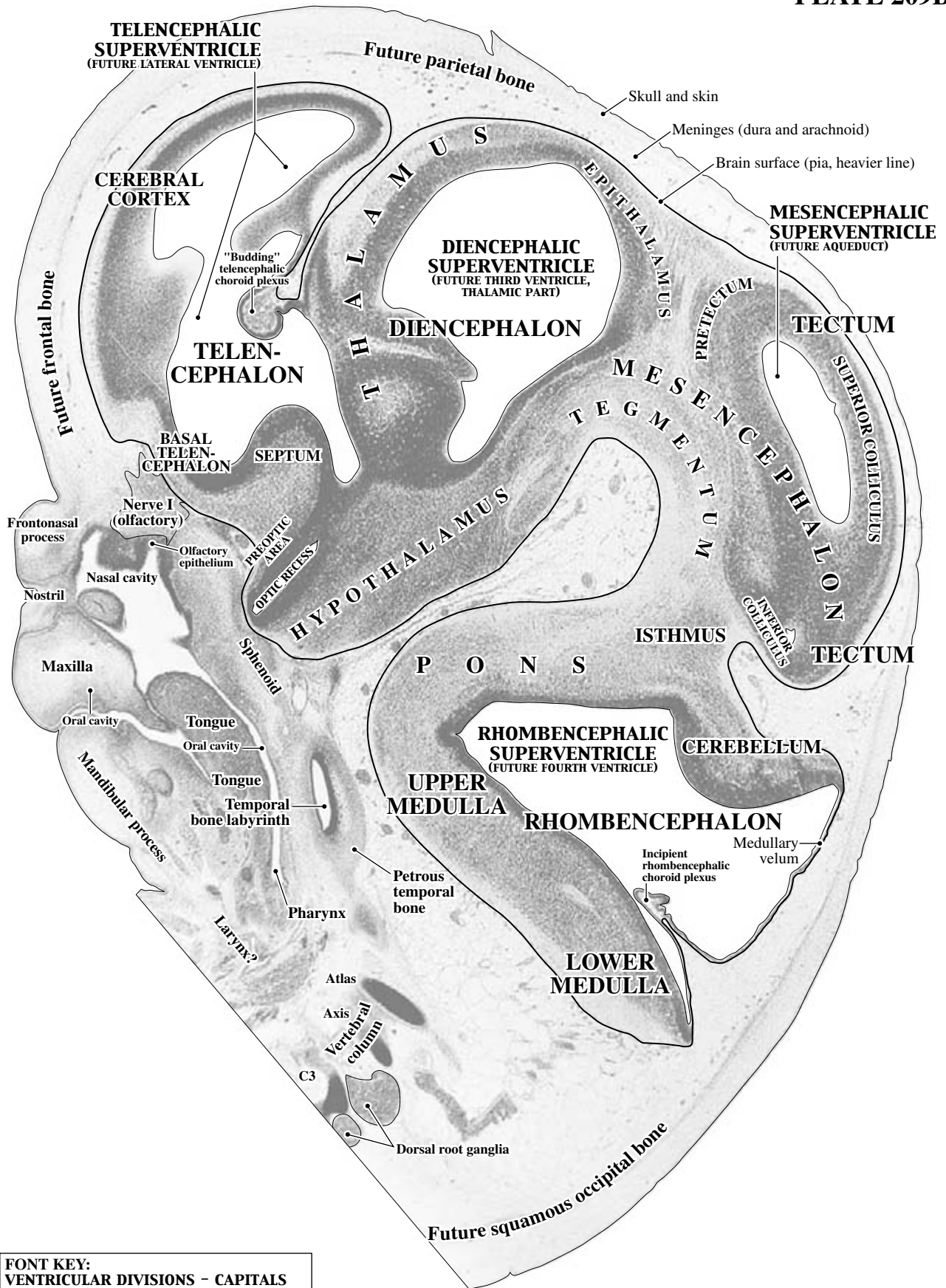
PLATE 209A

**GW7.5 Sagittal
CR 21 mm, C6202
Level 3: Slide 24, Section 2
Left side of brain
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



**Neuroepithelial and parenchymal
structures are labeled in Parts C and
D of this plate on the following pages.**

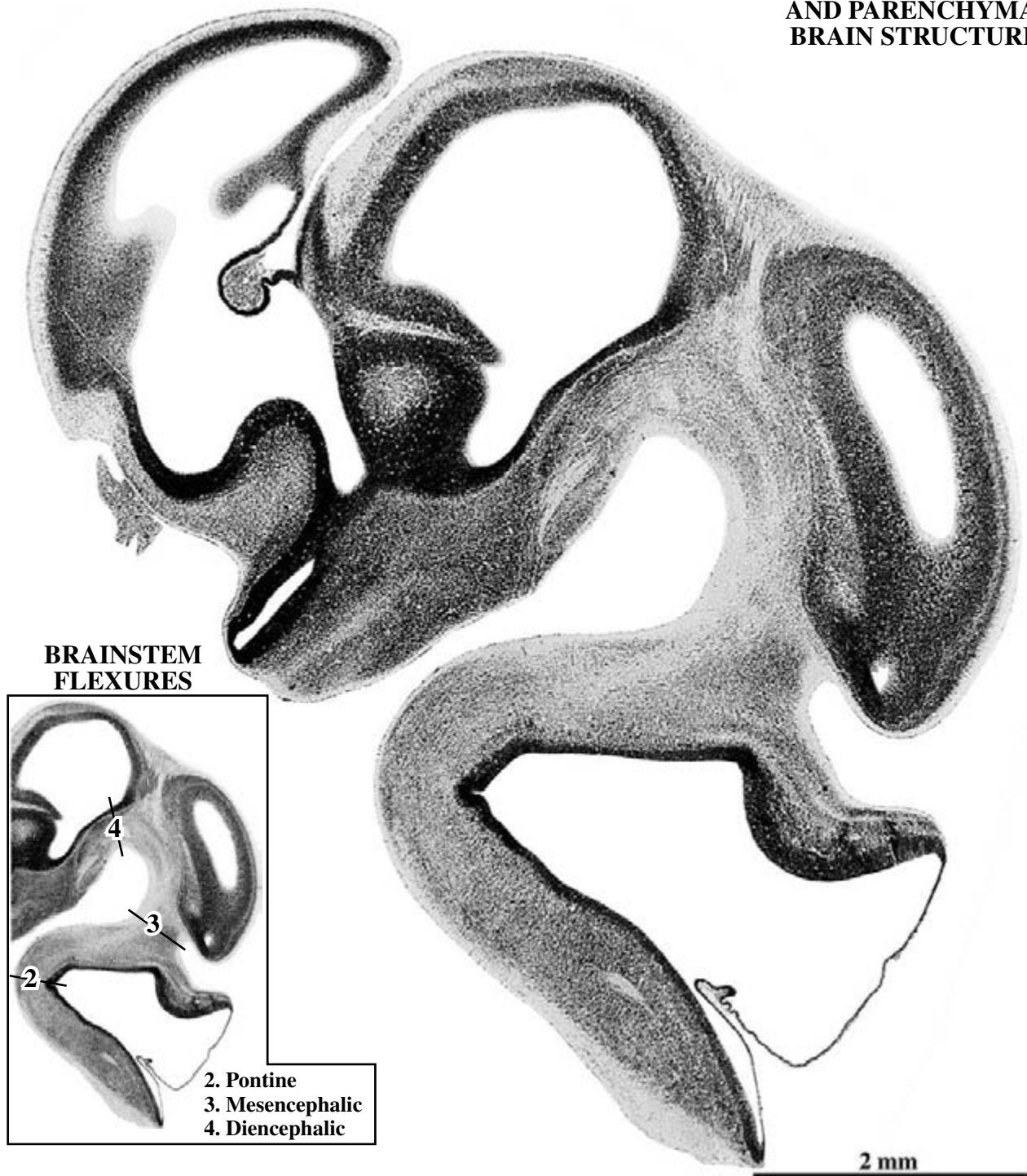
2 mm



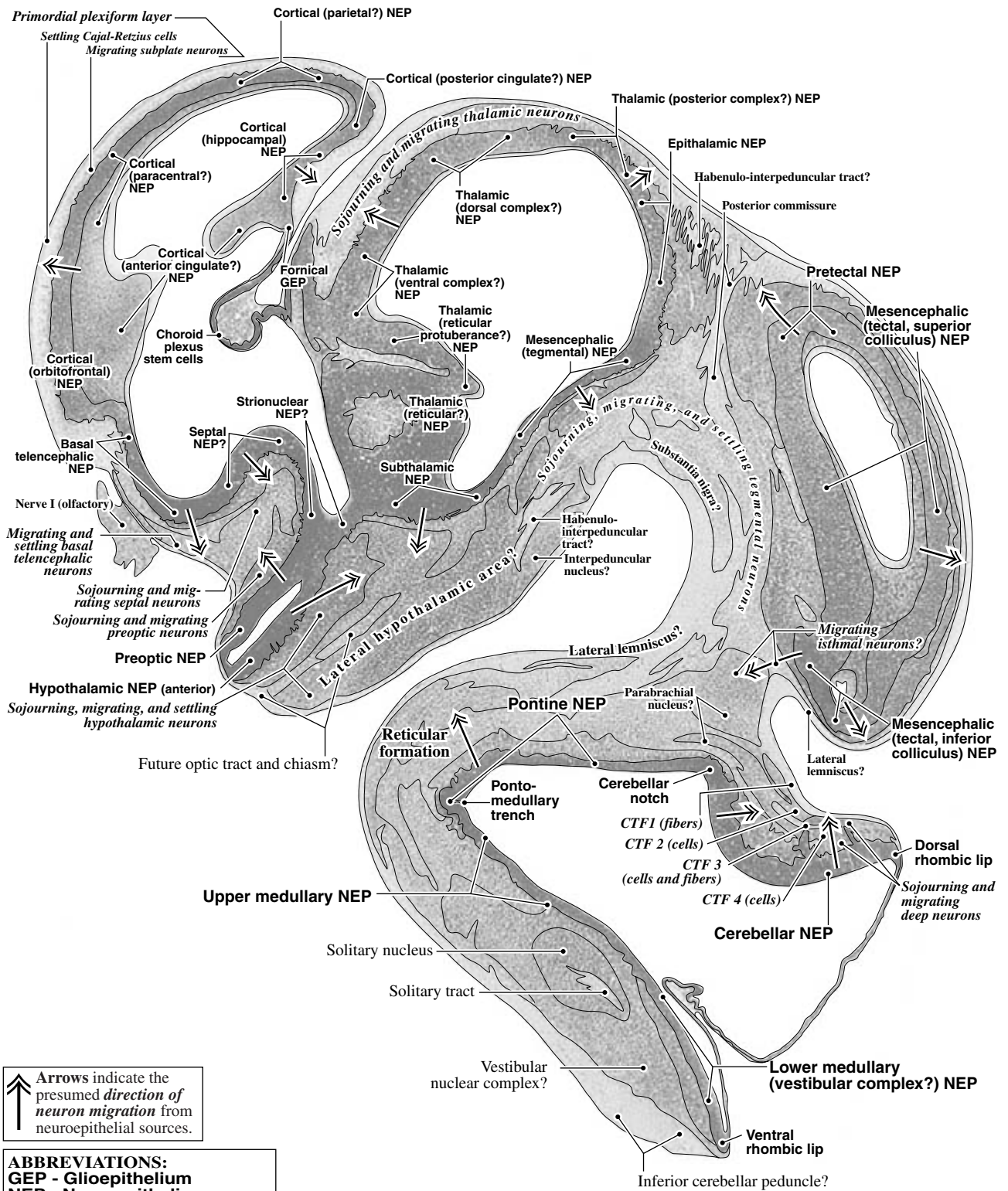
FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times Bold CAPITALS
 All other structures - Times Roman or Bold

PLATE 209C

GW7.5 Sagittal
CR 21 mm, C6202
Level 3: Slide 24, Section 2
Left side of brain
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES



The skull, major brain structures, and ventricular divisions are labeled in Parts A and B of this plate on the preceding pages.



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
CTF - Cerebellar transitional field

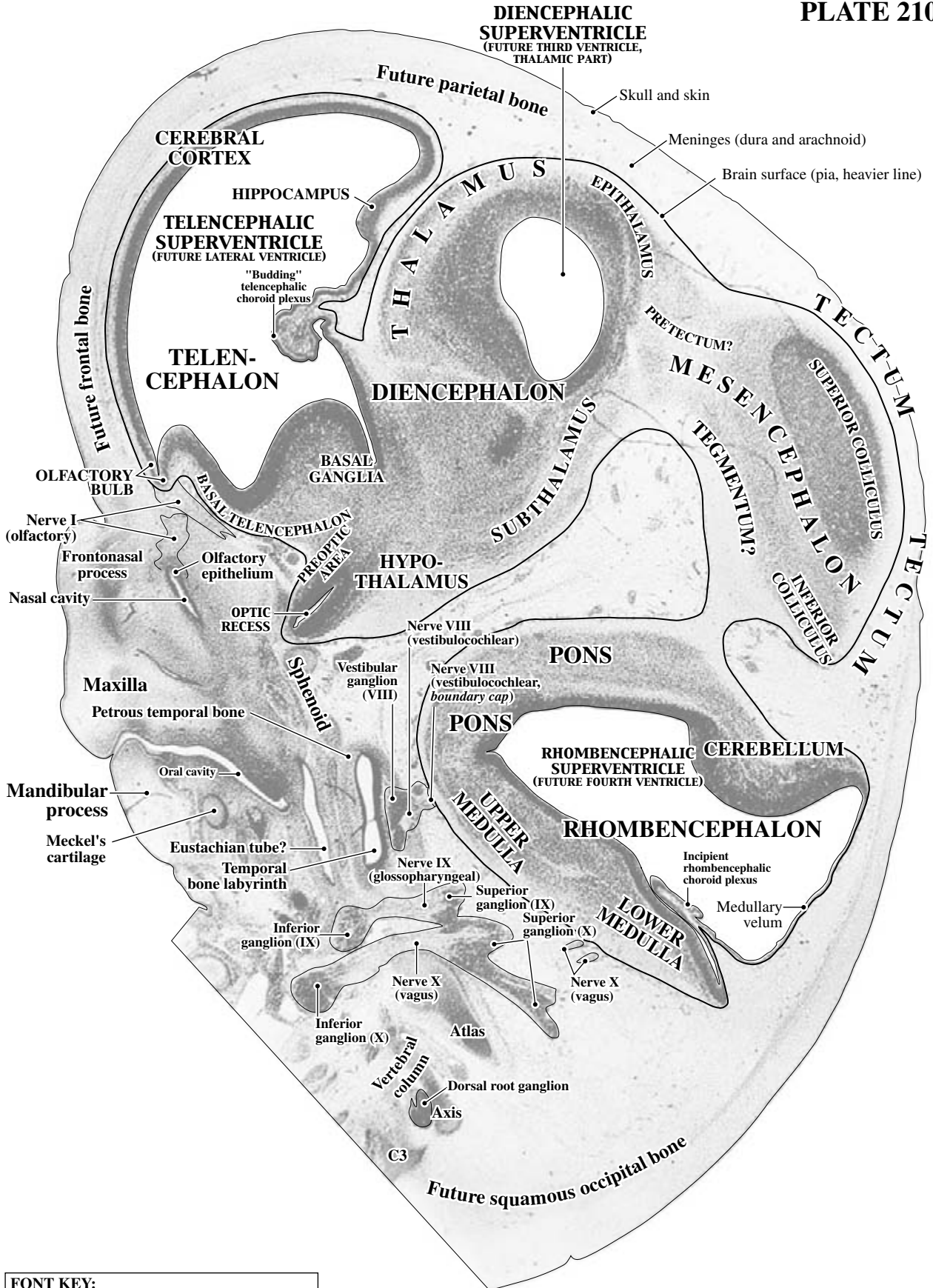
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Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

**GW7.5 Sagittal
CR 21 mm, C6202
Level 4: Slide 22, Section 2
Left side of brain
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



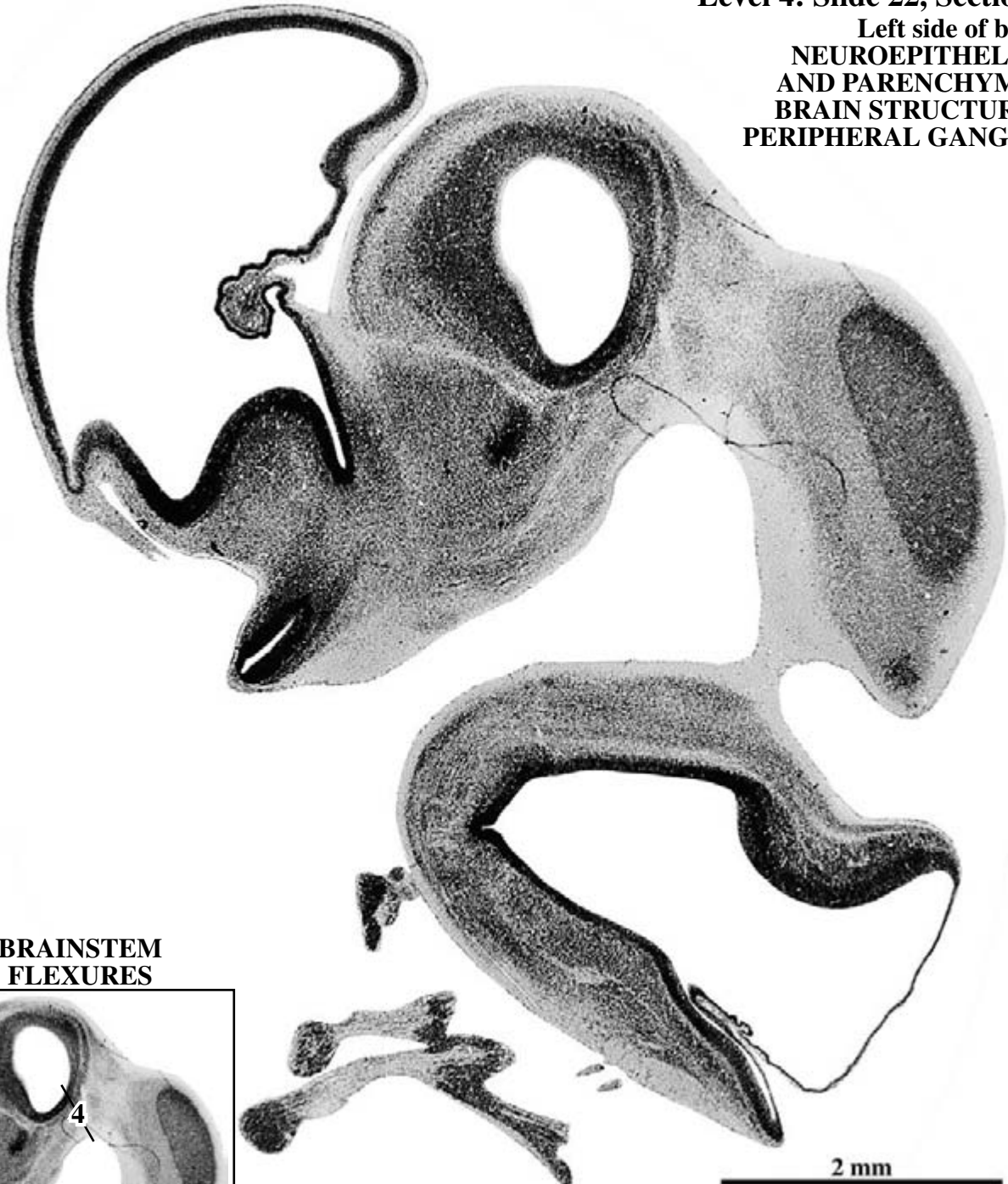
2 mm

**Neuroepithelial and parenchymal
structures are labeled in Parts C and
D of this plate on the following pages.**

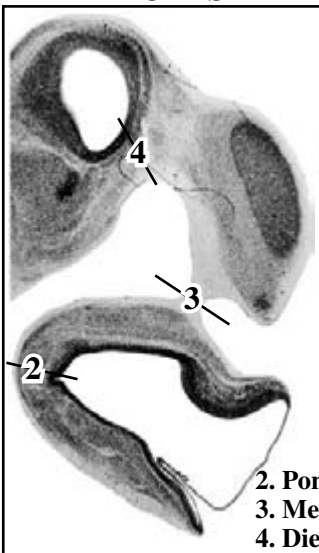


FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

GW7.5 Sagittal
CR 21 mm, C6202
Level 4: Slide 22, Section 2
Left side of brain
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES,
PERIPHERAL GANGLIA



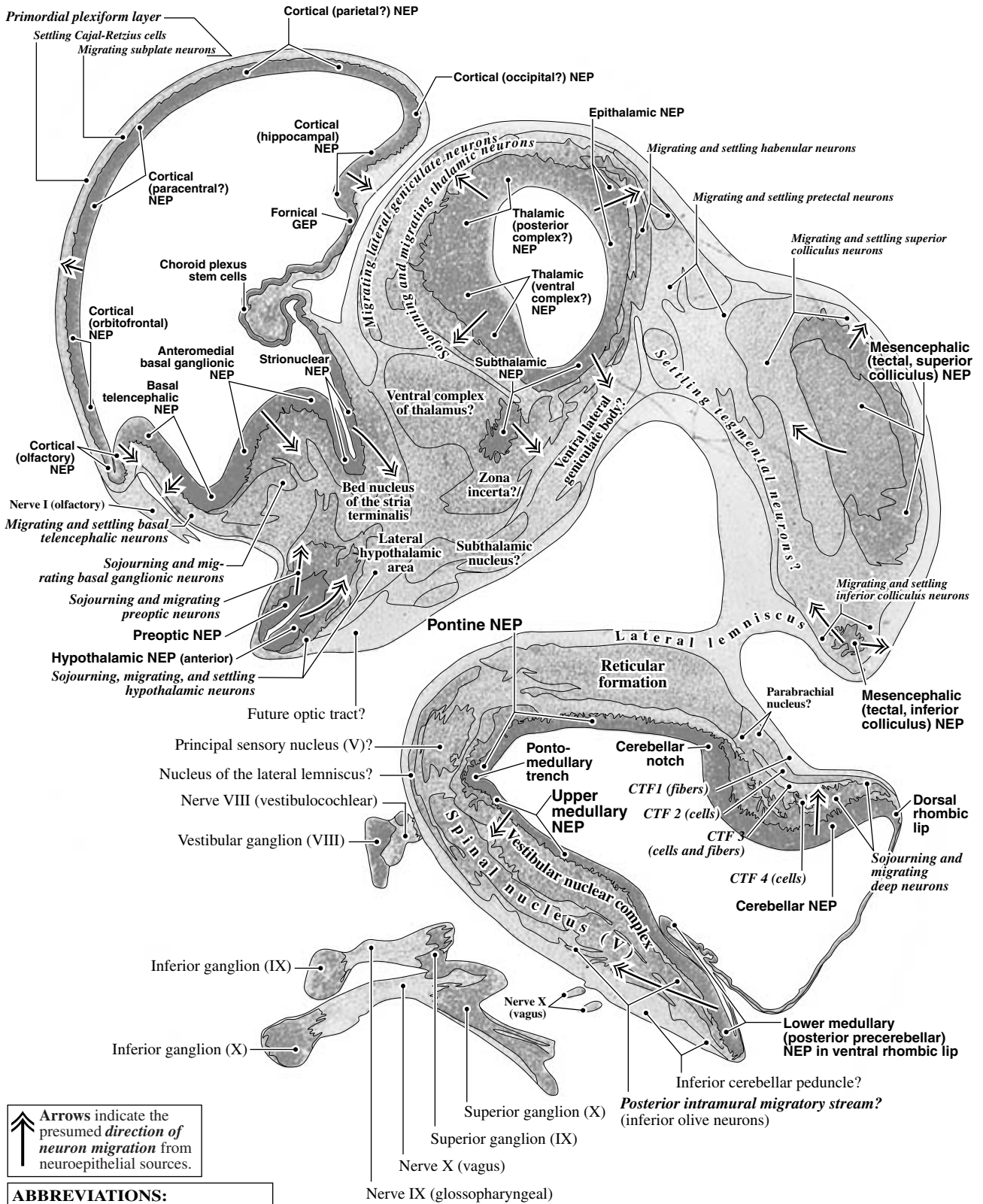
**BRAINSTEM
FLEXURES**



- 2. Pontine
- 3. Mesencephalic
- 4. Diencephalic

The skull, major brain structures,
and ventricular divisions are labeled
in Parts A and B of this plate on the
preceding pages.

PLATE 210D



↑ Arrows indicate the presumed direction of neuron migration from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
CTF - Cerebellar transitional field

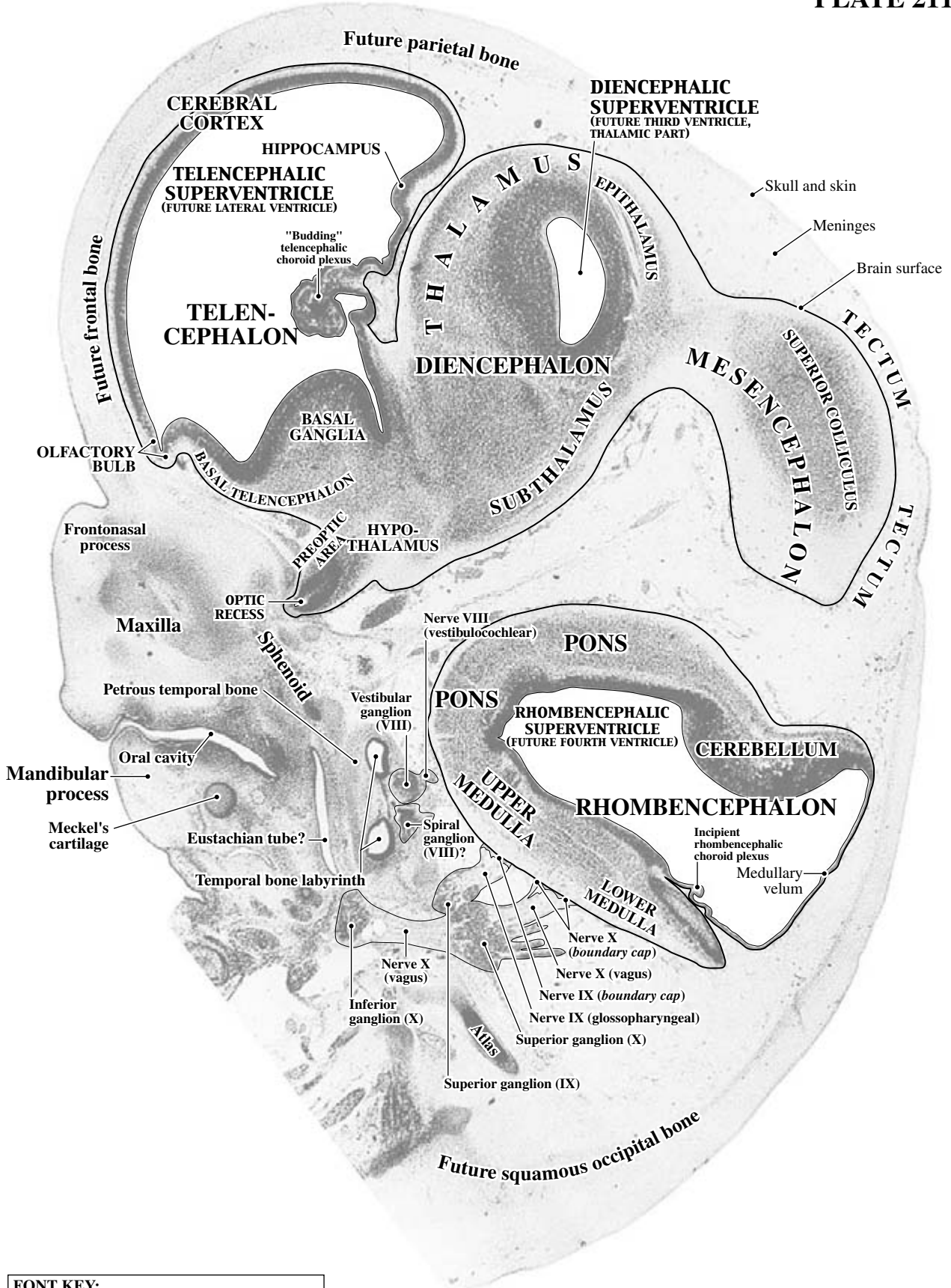
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Transient structure - Times bold italic
 Permanent structure - Times Roman or **Bold**

**GW7.5 Sagittal
CR 21 mm, C6202
Level 5: Slide 21, Section 2
Left side of brain
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



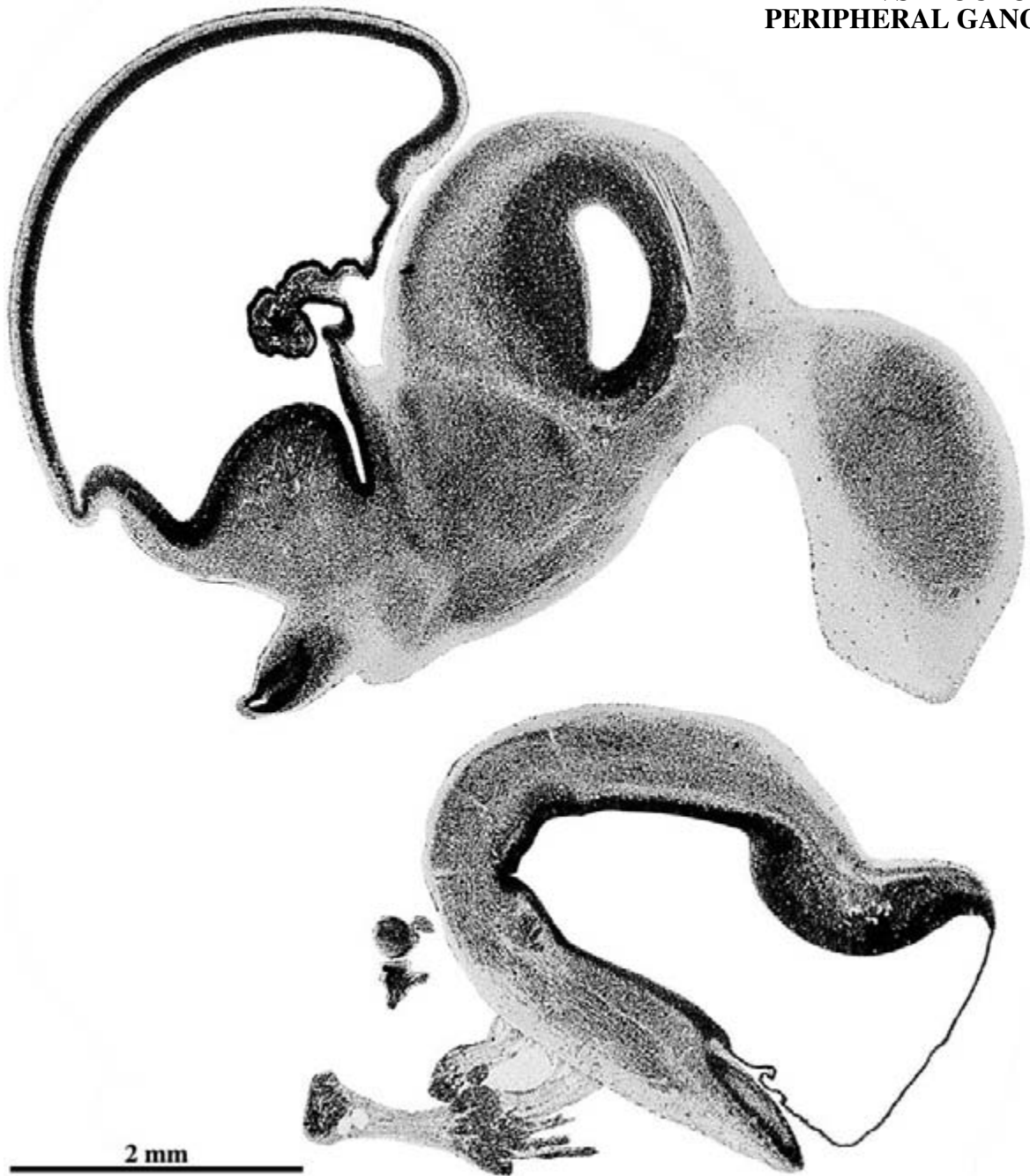
2 mm

**Neuroepithelial and parenchymal
structures are labeled in Parts C and
D of this plate on the following pages.**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

**GW7.5 Sagittal
CR 21 mm, C6202
Level 5: Slide 21, Section 2
Left side of brain
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES,
PERIPHERAL GANGLIA**



**The skull, major brain structures,
and ventricular divisions are
labeled in Parts A and B of this
plate on the preceding pages.**

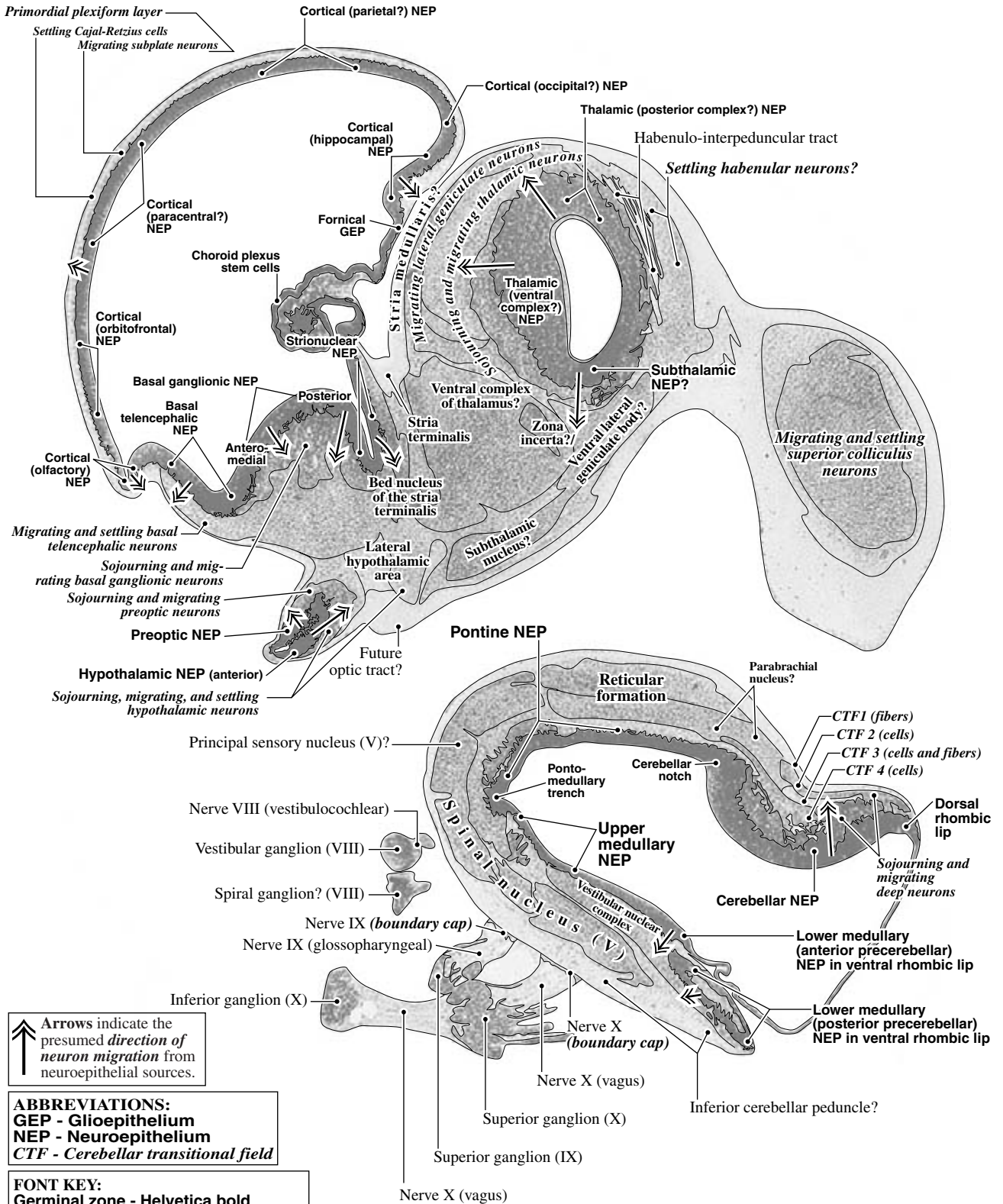


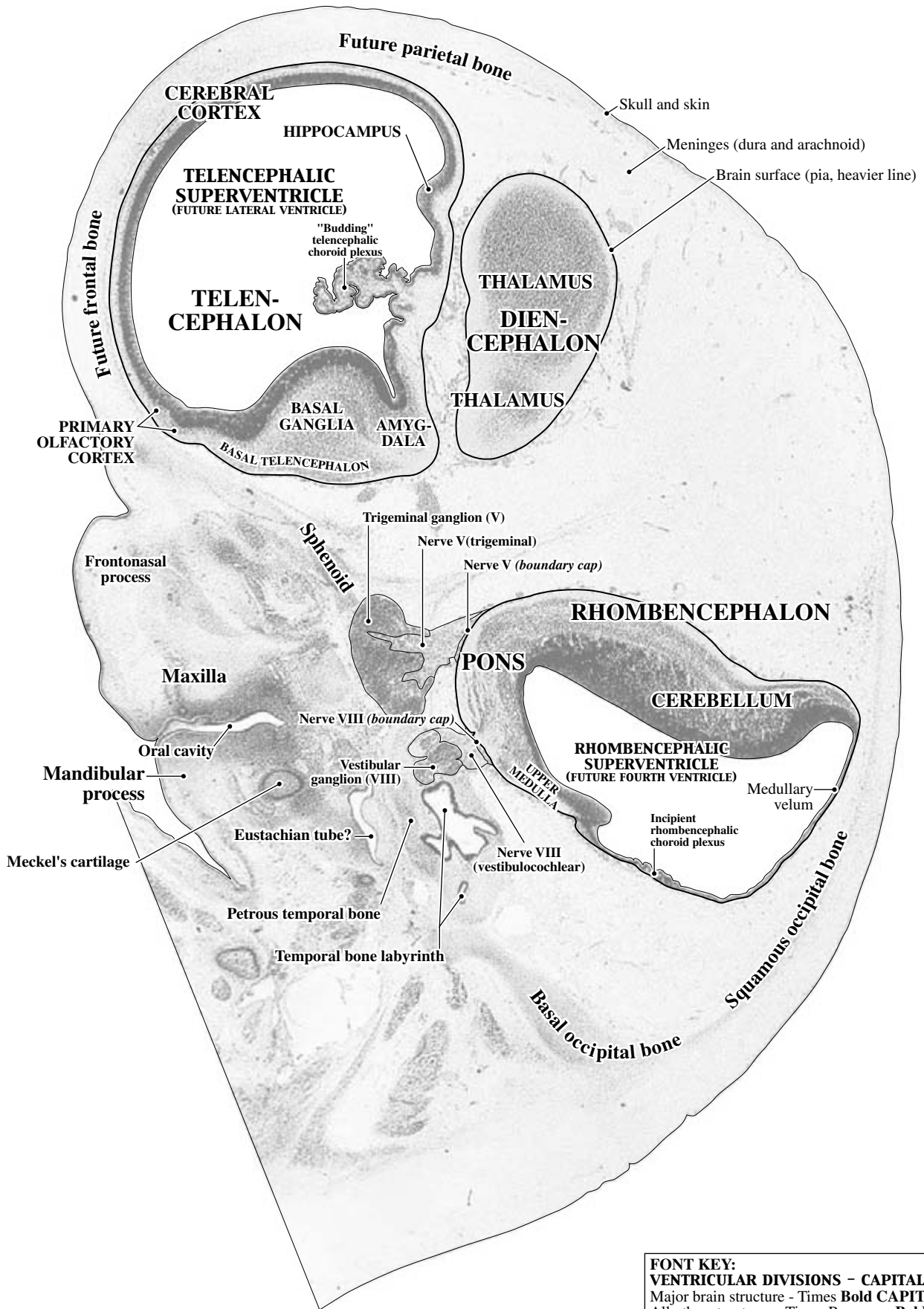
PLATE 212A

**GW7.5 Sagittal
CR 21 mm, C6202
Level 6: Slide 18, Section 2
Left side of brain
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



2 mm

**Neuroepithelial and parenchymal
structures are labeled in Parts C and
D of this plate on the following pages.**



FONT KEY:
VENTRICULAR DIVISIONS - CAPITALS
 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**

PLATE 212C

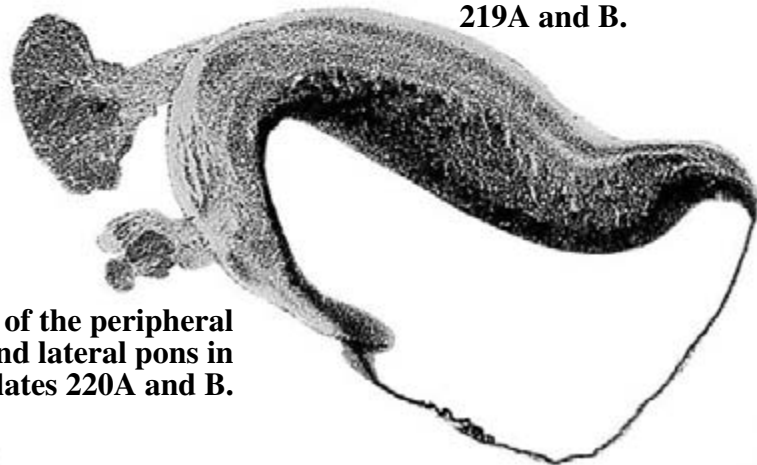
**GW7.5 Sagittal
CR 21 mm, C6202
Level 6: Slide 18, Section 2
Left side of brain
NEUROEPITHELIAL
AND PARENCHYMAL
BRAIN STRUCTURES,
PERIPHERAL GANGLIA**

See details of the cerebral
cortex in Plates 214A and B.



See details of the hippocampus,
basal ganglia, and amygdala in
Plates 216A and B.

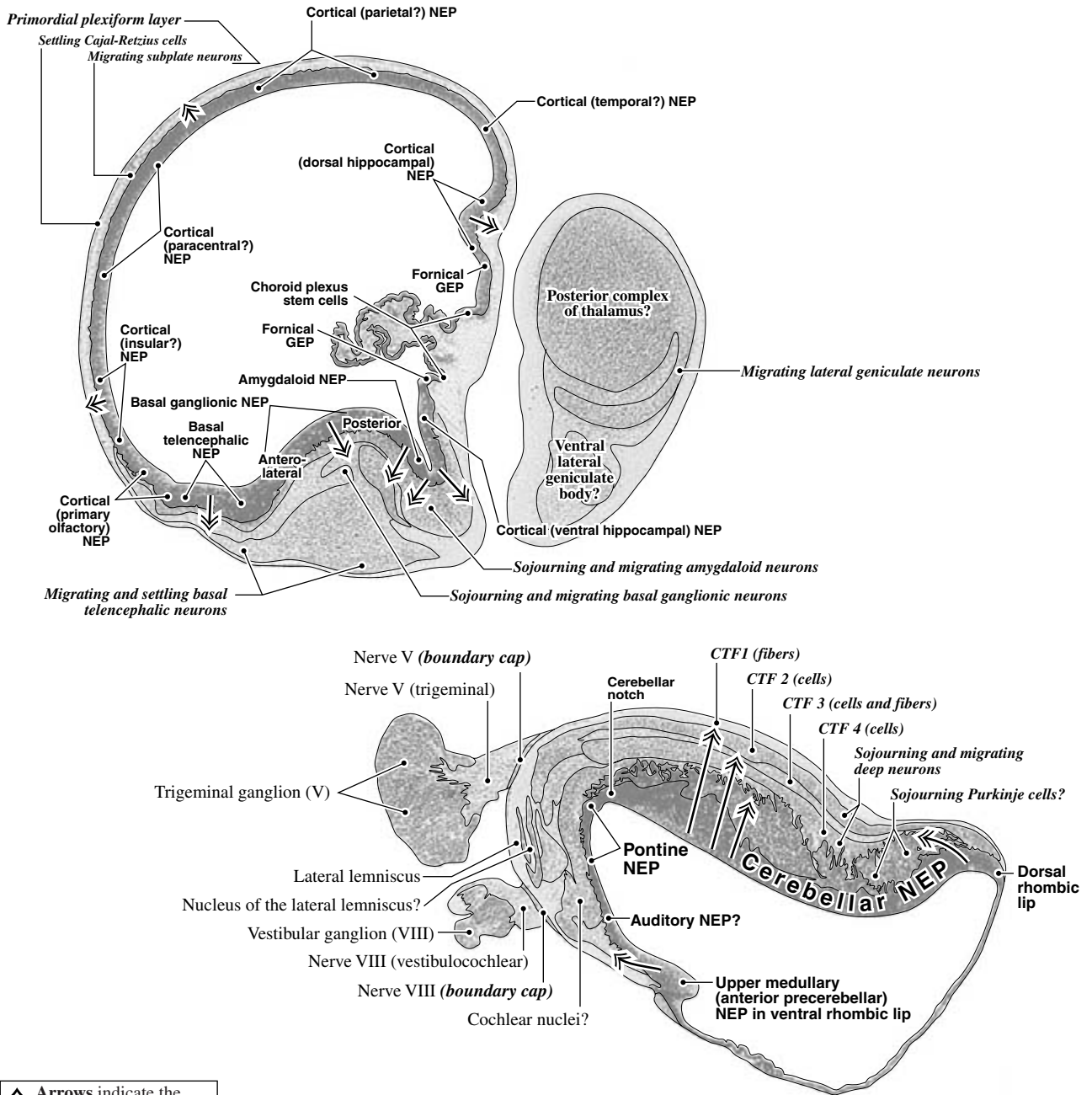
See details of the
cerebellum in Plates
219A and B.



See details of the peripheral
ganglia and lateral pons in
Plates 220A and B.

2 mm

The skull, major brain structures,
and ventricular divisions are
labeled in Parts A and B of this
plate on the preceding pages.



↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
CTF - Cerebellar transitional field

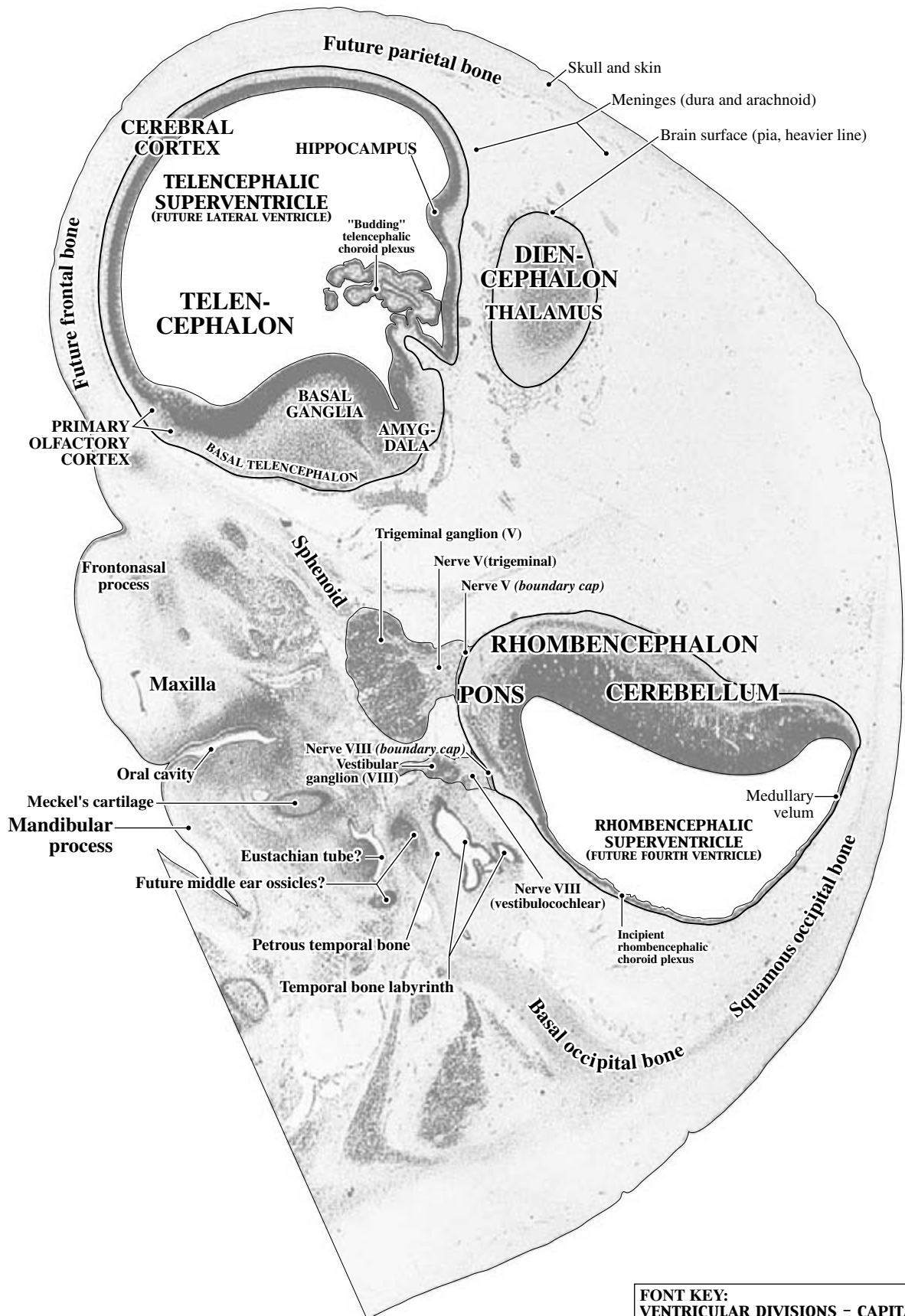
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Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 213A

**GW7.5 Sagittal
CR 21 mm, C6202
Level 7: Slide 17, Section 2
Left side of brain
SKULL, MAJOR BRAIN
STRUCTURES, AND
VENTRICULAR
DIVISIONS**



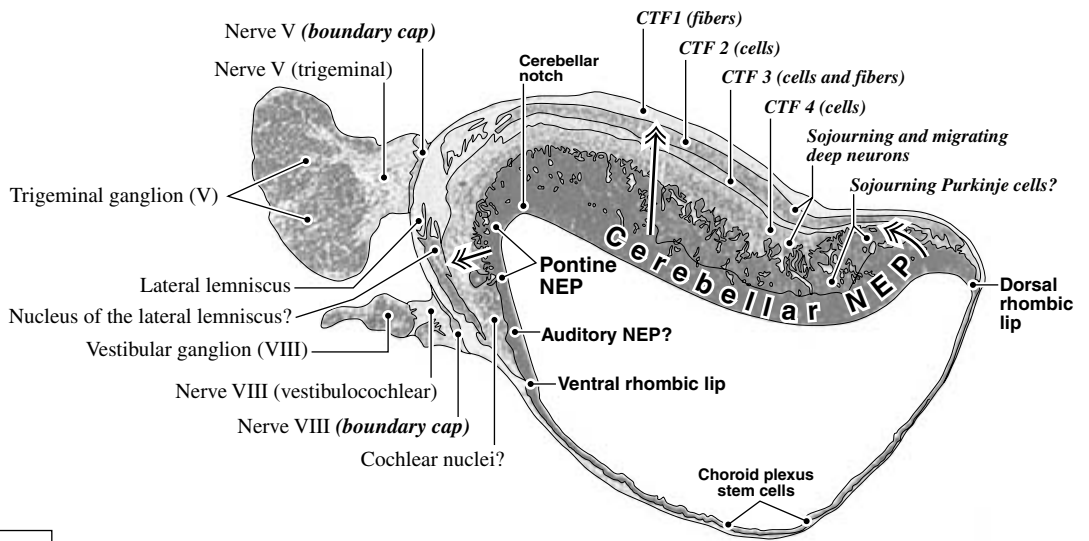
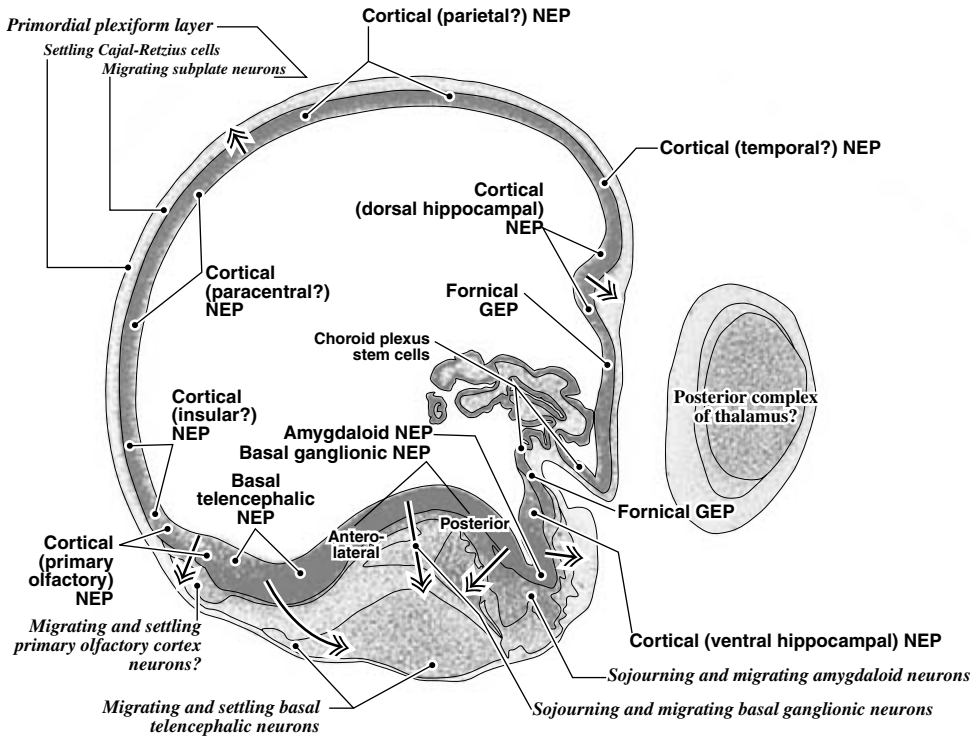
**Neuroepithelial and parenchymal
structures are labeled in Parts C and
D of this plate on the following pages.**



FONT KEY:
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 Major brain structure - Times **Bold CAPITALS**
 All other structures - Times Roman or **Bold**



The skull, major brain structures,
and ventricular divisions are
labeled in Parts A and B of this
plate on the preceding pages.



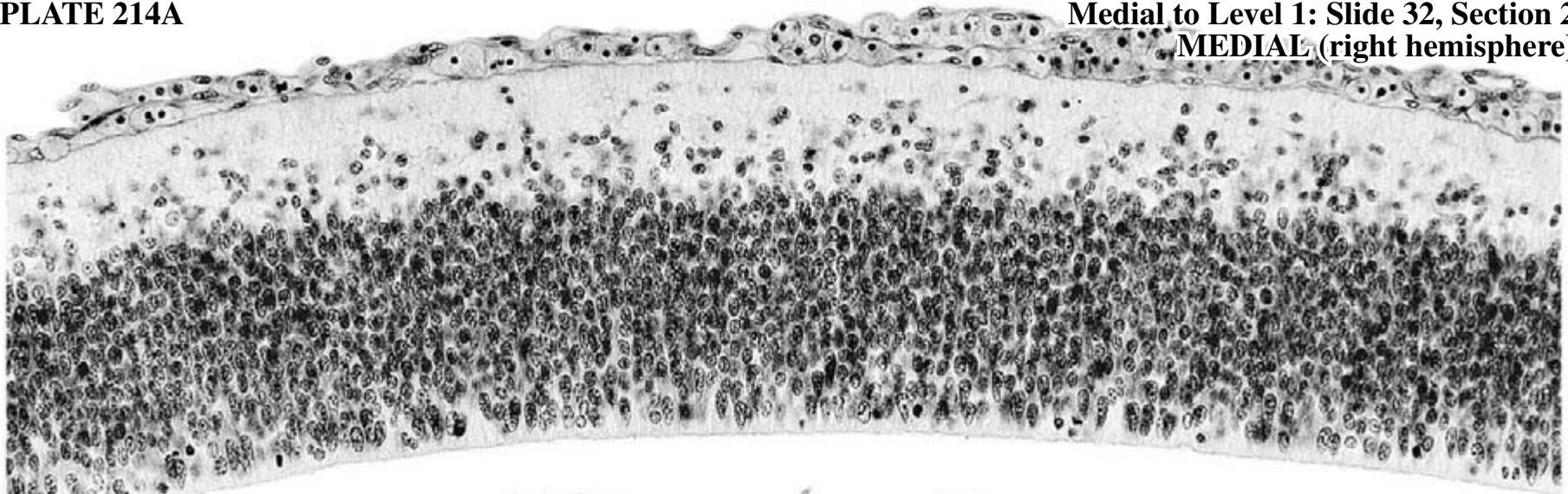
↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

ABBREVIATIONS:
GEP - Glioepithelium
NEP - Neuroepithelium
CTF - Cerebellar transitional field

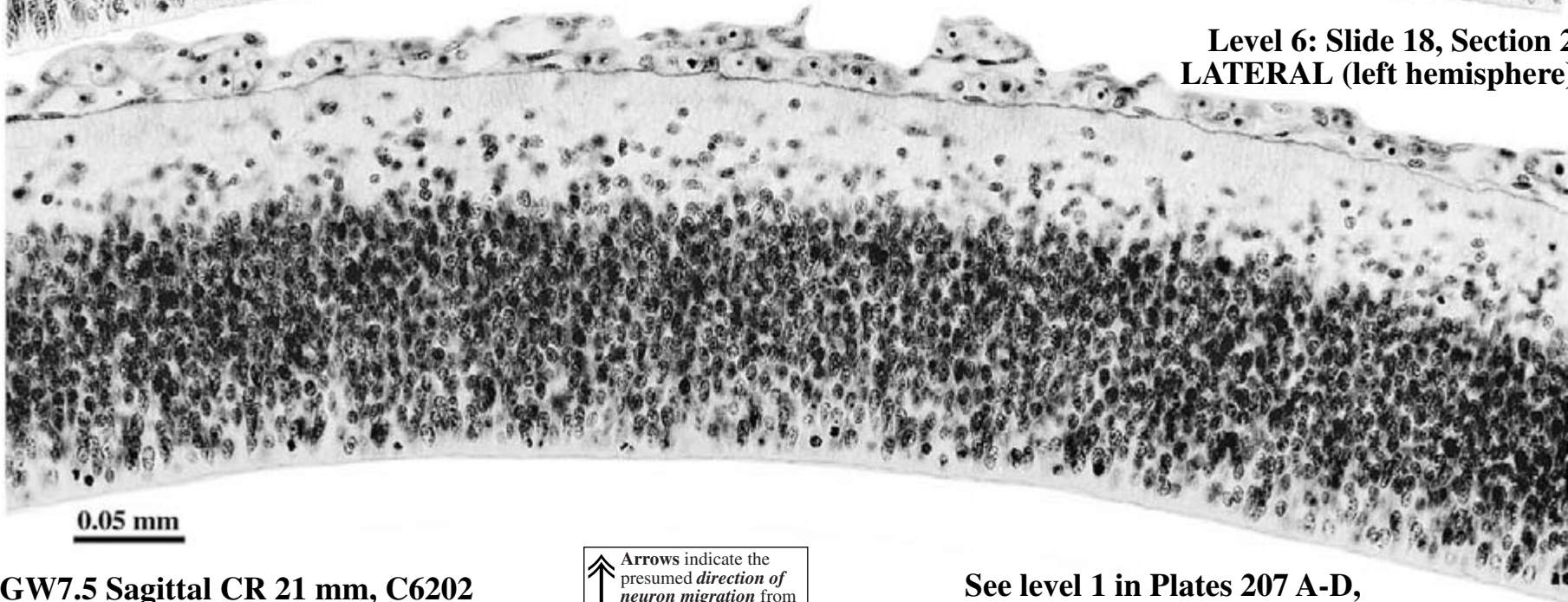
FONT KEY:
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Transient structure - Times bold italic
Permanent structure - Times Roman or Bold

PLATE 214A

**Medial to Level 1: Slide 32, Section 2
MEDIAL (right hemisphere)**



**Level 6: Slide 18, Section 2
LATERAL (left hemisphere)**



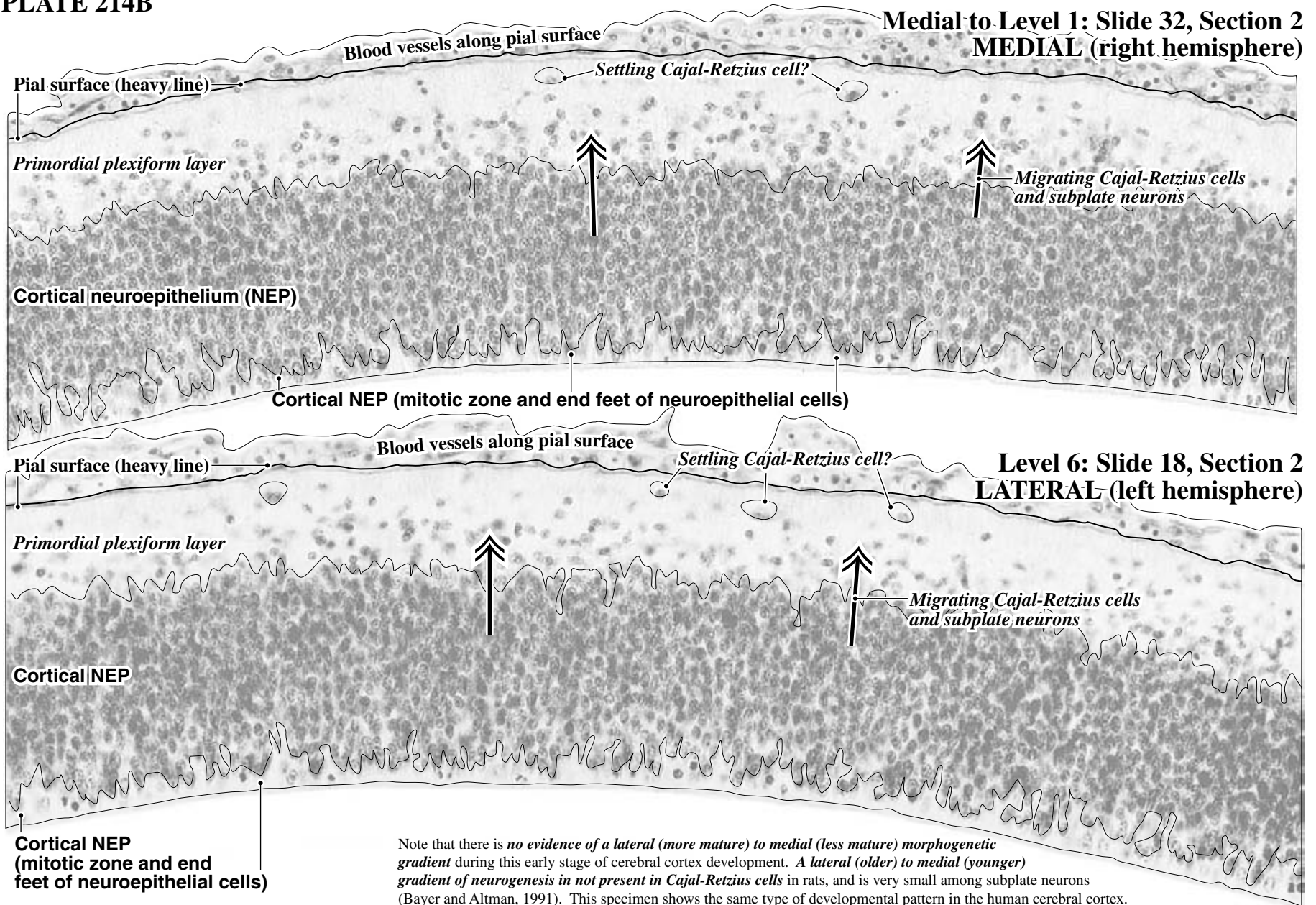
0.05 mm

**GW7.5 Sagittal CR 21 mm, C6202
DORSAL CEREBRAL CORTEX**

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

**See level 1 in Plates 207 A-D,
level 6 in Plates 212 A-D.**

PLATE 214B



Note that there is *no evidence of a lateral (more mature) to medial (less mature) morphogenetic gradient* during this early stage of cerebral cortex development. A *lateral (older) to medial (younger) gradient of neurogenesis is not present in Cajal-Retzius cells* in rats, and is very small among subplate neurons (Bayer and Altman, 1991). This specimen shows the same type of developmental pattern in the human cerebral cortex.

PLATE 215A

**GW7.5 Sagittal
CR 21 mm, C6202
Medial to Level 1:
Slide 31, Section 2
Right side of brain
OLFACTORY BULB
AND BASAL
TELENCEPHALON**

**See level 1 in
Plates 207A-D.**

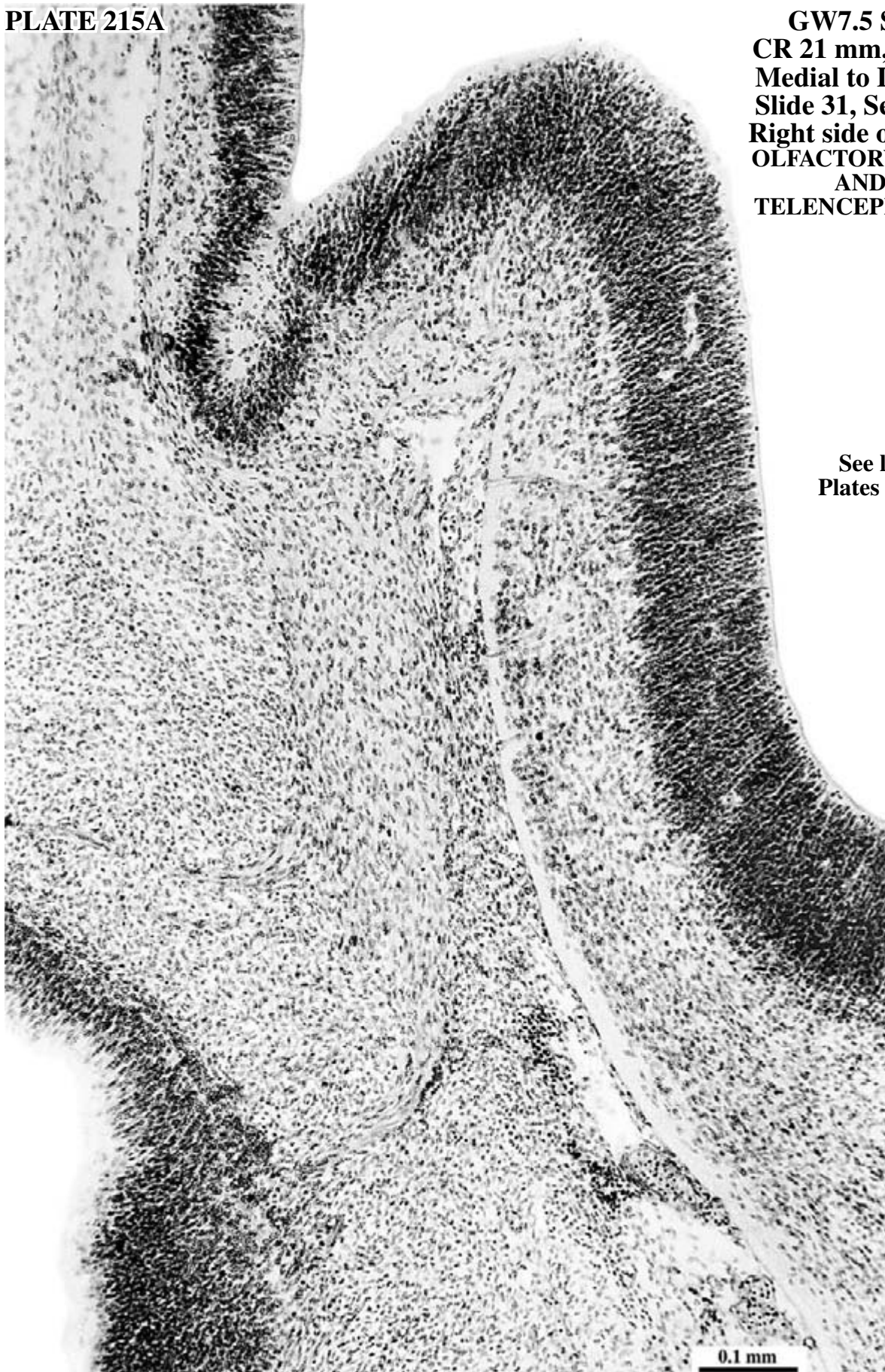


PLATE 215B

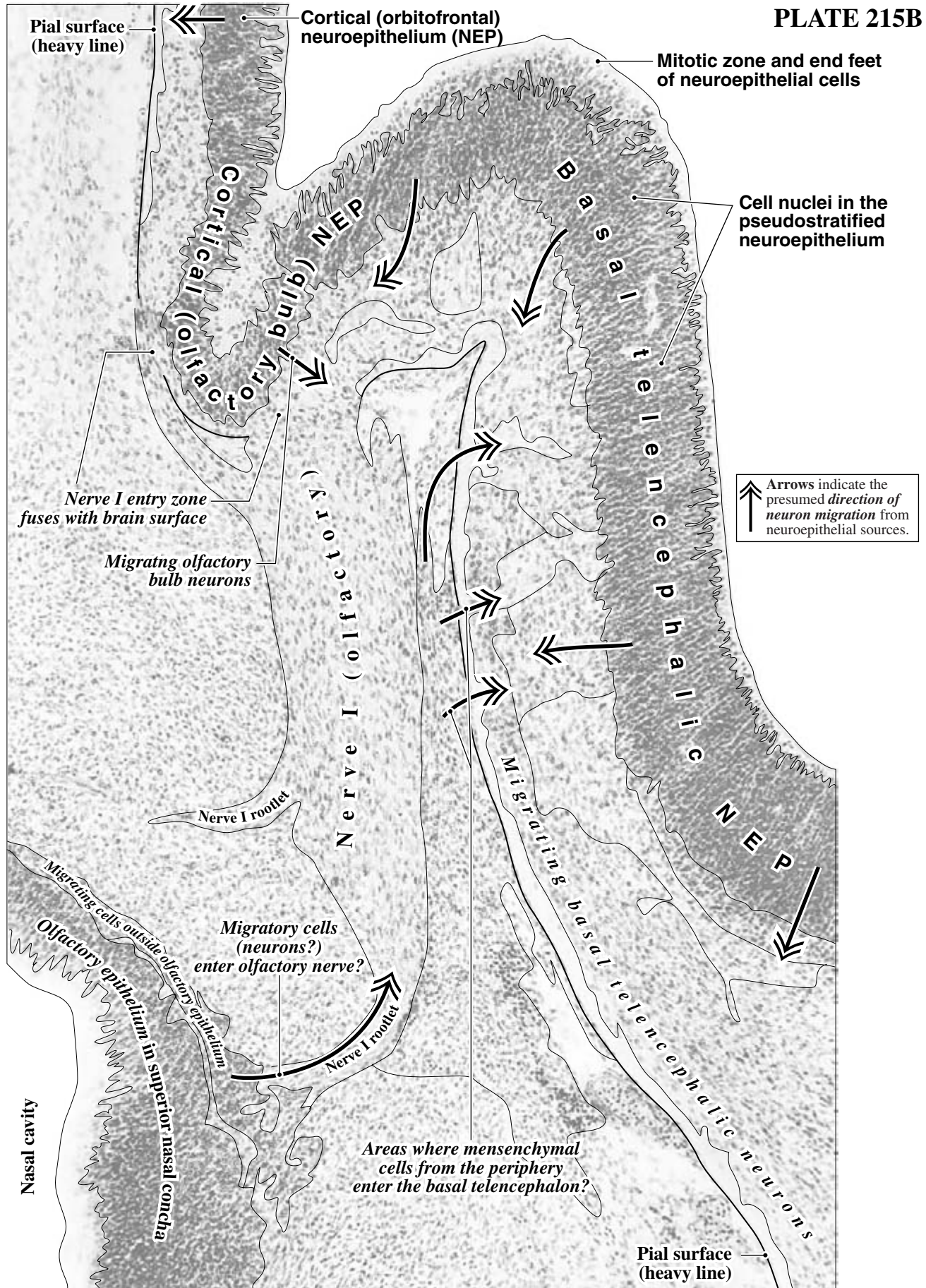
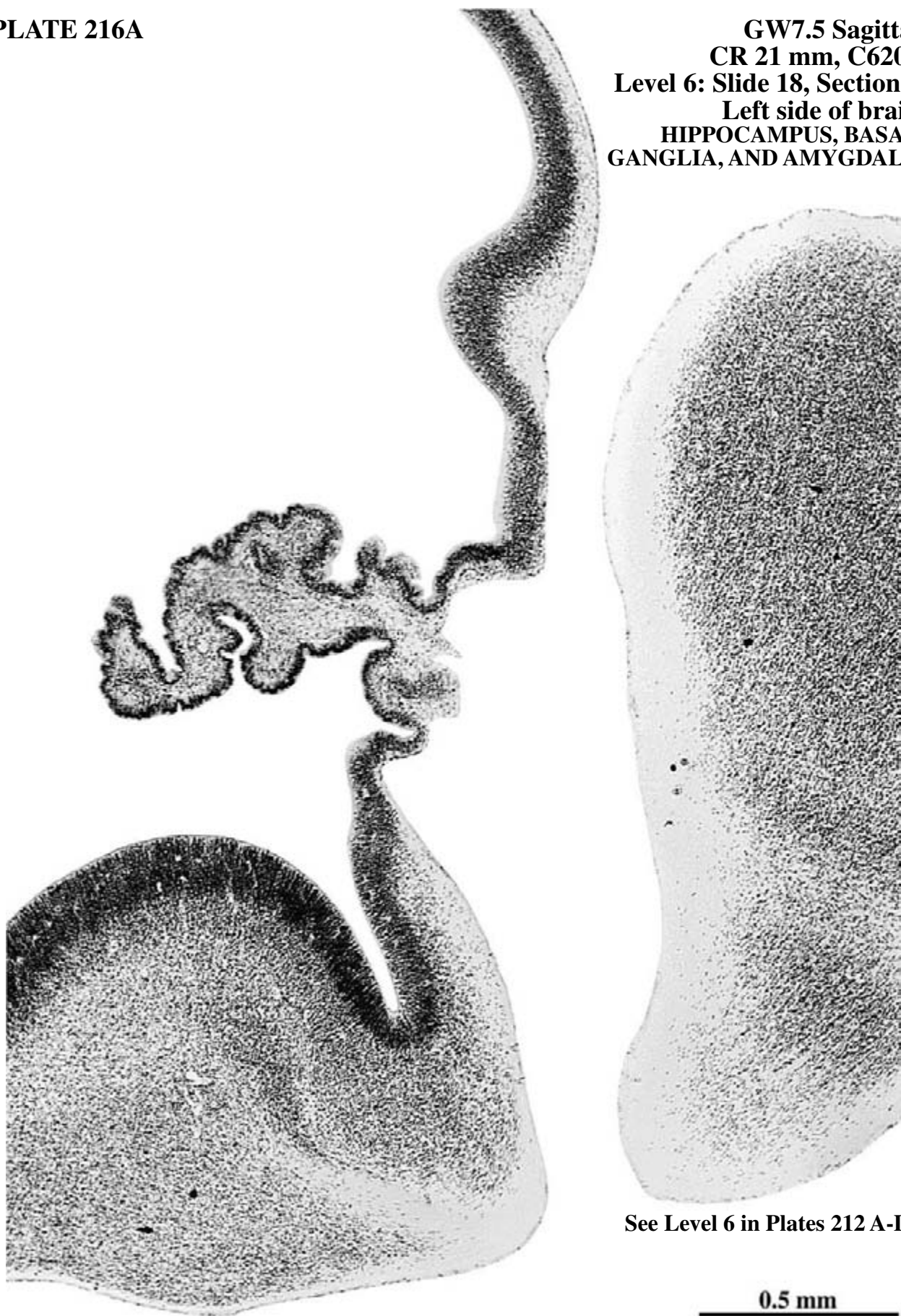


PLATE 216A

**GW7.5 Sagittal
CR 21 mm, C6202
Level 6: Slide 18, Section 2
Left side of brain
HIPPOCAMPUS, BASAL
GANGLIA, AND AMYGDALA**



See Level 6 in Plates 212 A-D.

0.5 mm

TELENCEPHALON

TELENCEPHALIC
SUPERVENTRICLE
(FUTURE LATERAL VENTRICLE)

Primordial plexiform layer

Settling Cajal-Retzius cells

Migrating subplate neurons

Cortical
(temporal?) NEP

Arrows indicate the
presumed direction of
neuron migration from
neuroepithelial sources.

Cortical (dorsal hippocampal) NEP

DORSAL
HIPPOCAMPUS

Fornical GEP

Choroid plexus stem cells

Vascular bed
(stroma)

"Budding"
telencephalic
choroid plexus

Choroid plexus
stem cells

DIENCEPHALON

THALAMUS

Posterior complex?

Basal ganglionic
NEP

Fornical
GEP?

VENTRAL
HIPPOCAMPUS?

Cortical
(ventral
hippocampal?)
NEP

Posterior
Antero-lateral
Sojourning and migrating
basal ganglionic neurons
BASAL GANGLIA
(putamen and globus pallidus?)

Amygdaloid
NEP

AMYGDALA

THALAMUS

Ventral lateral
geniculate body?

Migrating and settling basal
telencephalic neurons

BASAL
TELENCEPHALON

Ansa lenticularis?

Sojourning and migrating
amygdaloid neurons

ABBREVIATIONS:
GEP - Glioeepithelium
NEP - Neuroepithelium

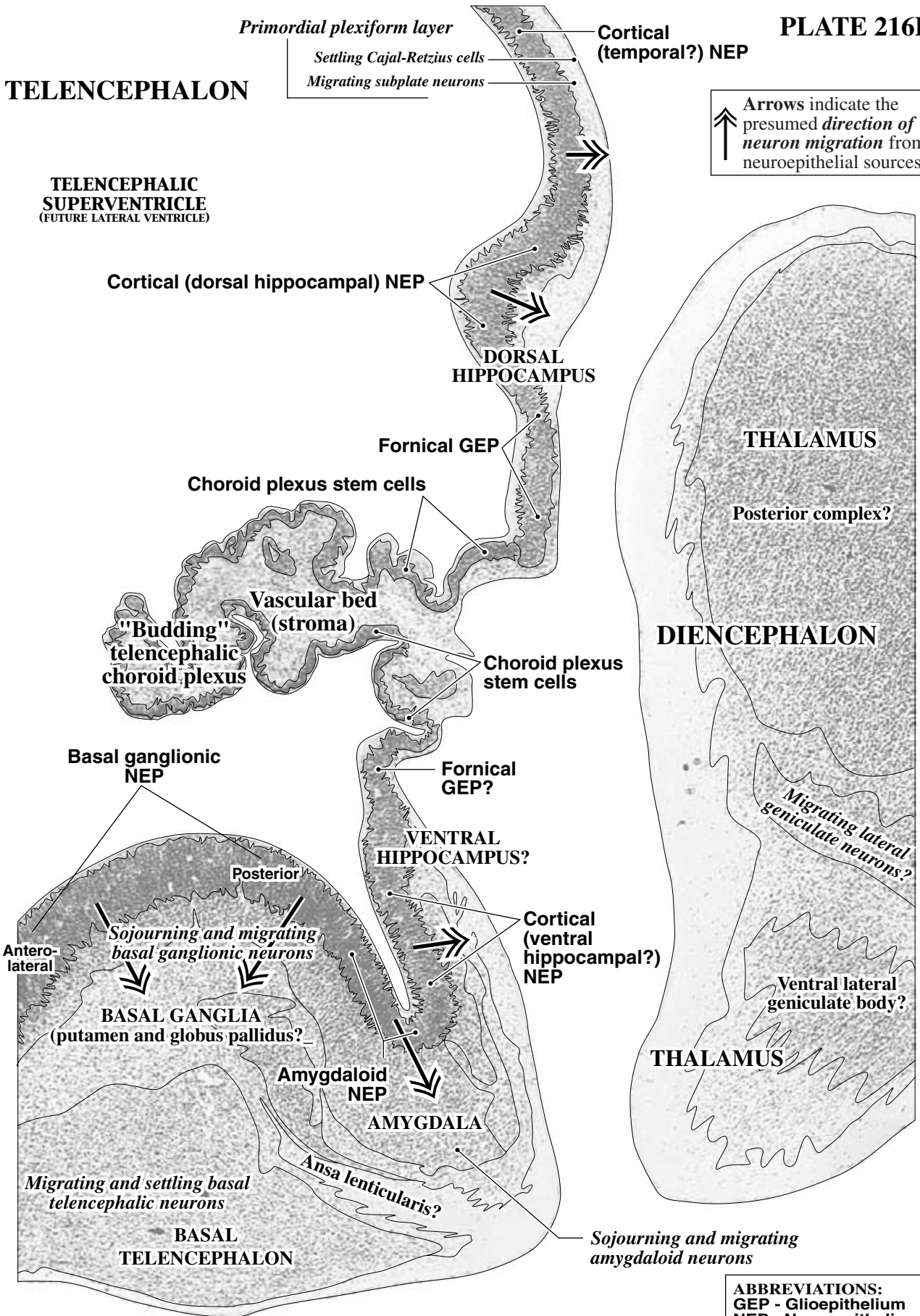


PLATE 217A

**GW7.5 Sagittal CR 21 mm, C6202
Between Levels 1 and 2: Slide 28, Section 3
HYPOTHALAMUS**



See level 1 in Plates 207A-D, level 2 in Plates 208 A-D.

PLATE 217B

Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

**DIENCEPHALIC SUPERVENTRICLE
(FUTURE THIRD VENTRICLE, HYPOTHALAMIC PART)**

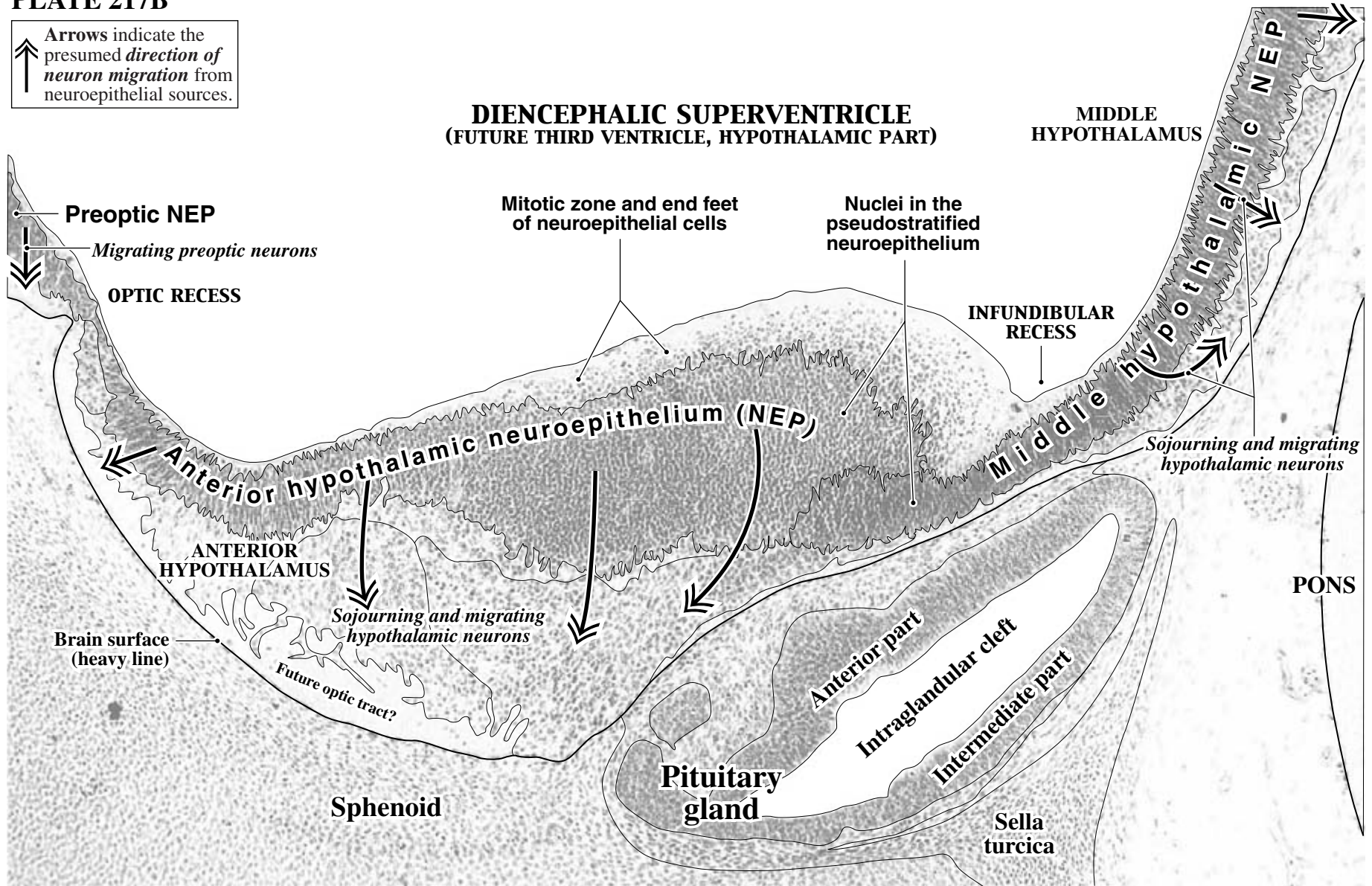
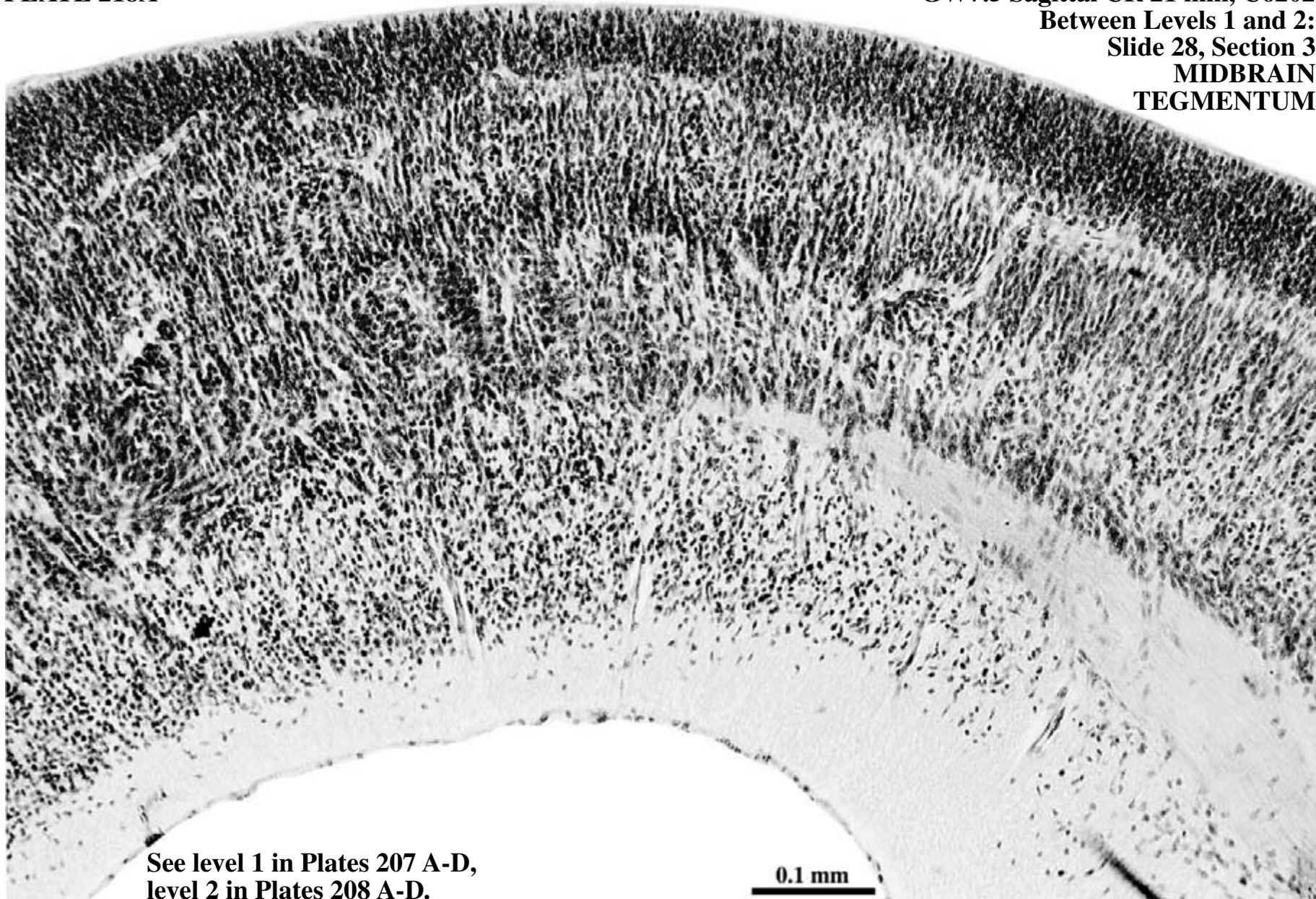


PLATE 218A

**GW7.5 Sagittal CR 21 mm, C6202
Between Levels 1 and 2:
Slide 28, Section 3
MIDBRAIN
TEGMENTUM**

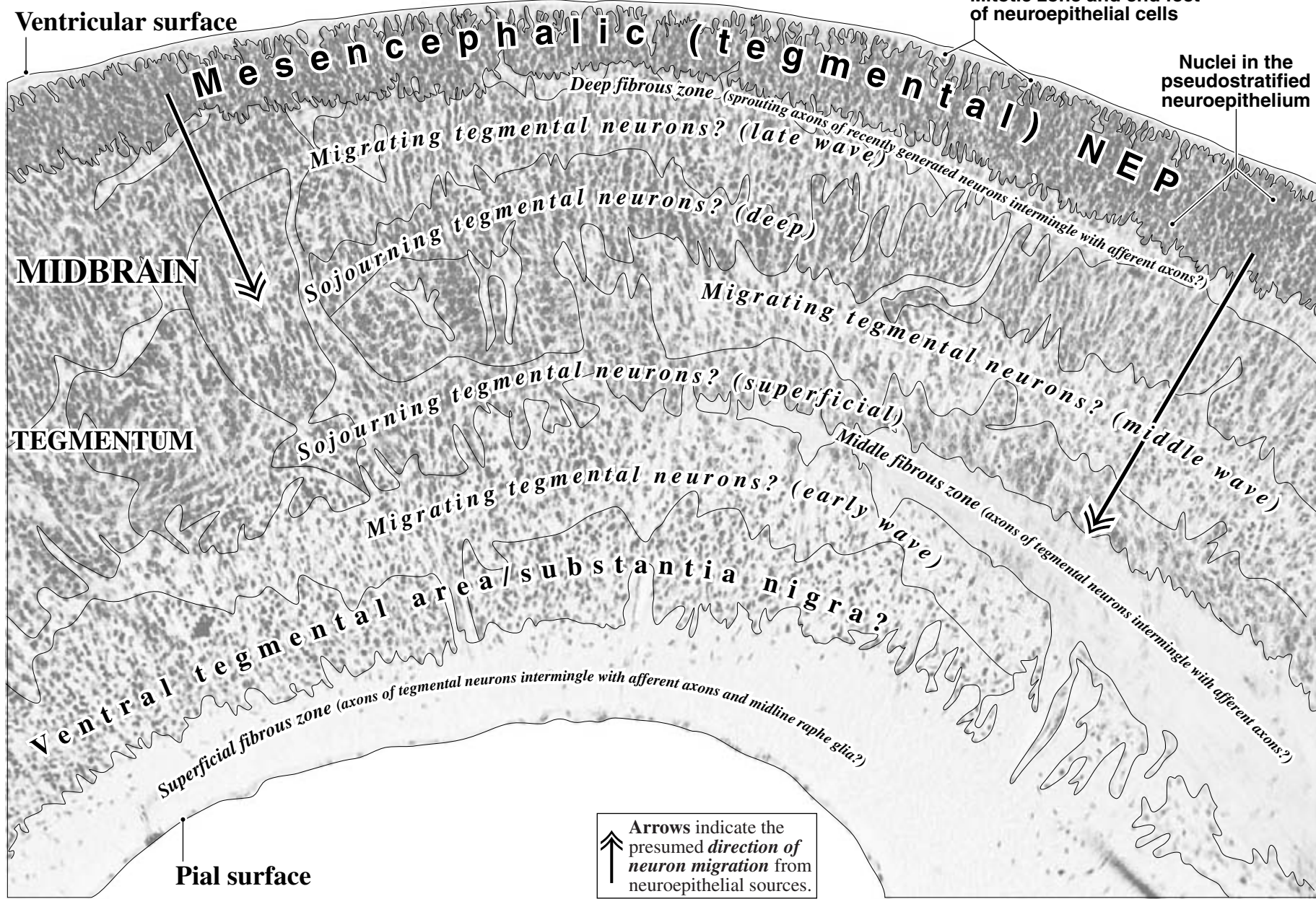


**See level 1 in Plates 207 A-D,
level 2 in Plates 208 A-D.**

0.1 mm

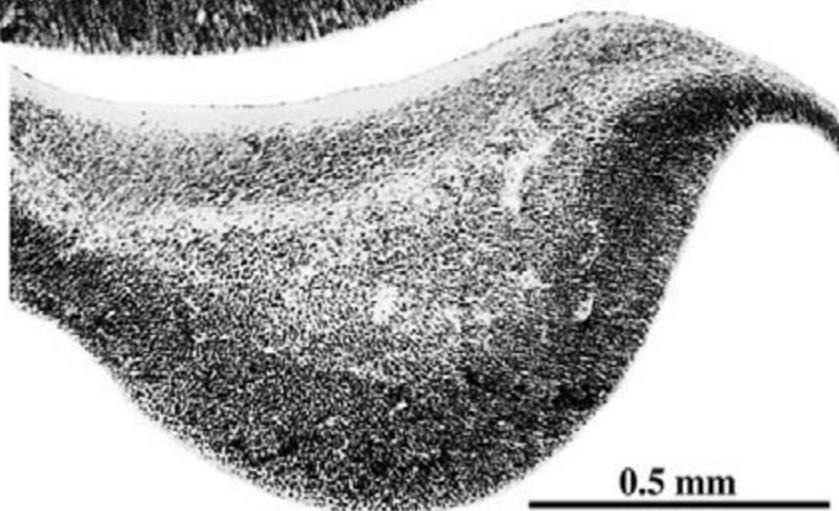
PLATE 218B

MESENCEPHALIC SUPERVENTRICLE (FUTURE AQUEDUCT)





HEMISPHERE
Level 6: Slide 18, Section 2



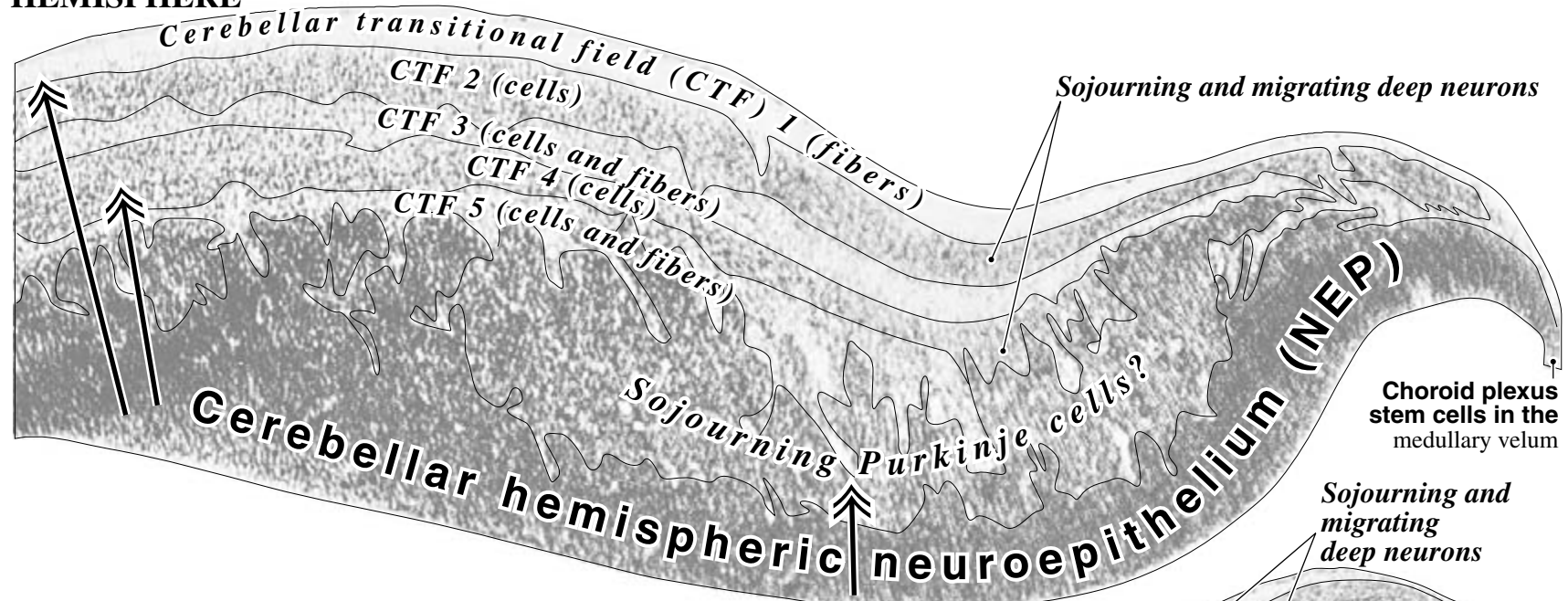
VERMIS
Between Levels 1 and 2:
Slide 29, Section 2

0.5 mm

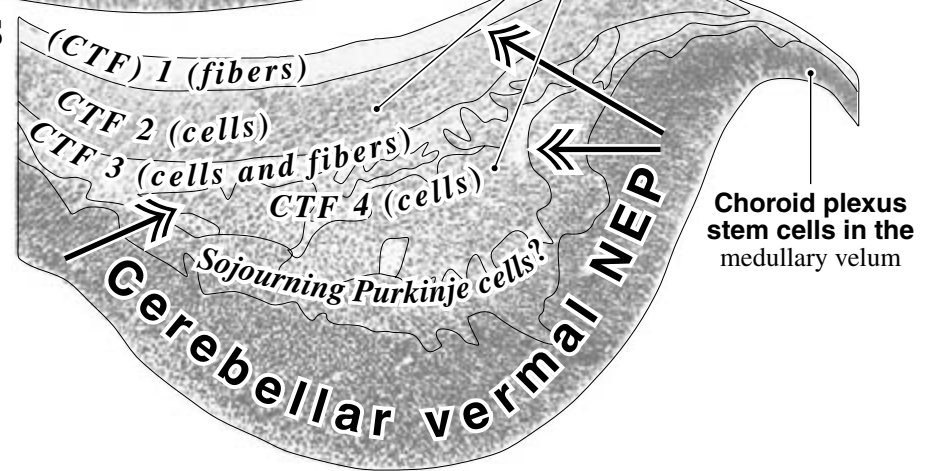
See level 1 in Plates 207 A-D, level 2 in Plates 208 A-D, and level 6 in Plates 212 A-D.

PLATE 219B

HEMISPHERE



VERMIS



Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

There is a lateral (more mature) to medial (less mature) morphogenetic gradient between the hemisphere and vermis of the cerebellum. The larger hemisphere has better defined and more transitional layers than the smaller vermis. Deep neurons predominate in the transitional field outside the germinal matrix throughout the cerebellum, indicating that they are earlier-generated neurons. Within both the hemispheric and vermal parts of the **cerebellar neuroepithelium**, more Purkinje cells and fewer deep neurons are being generated. Note that the **external germinal layer** is not present at this stage of cerebellar development.

PLATE 220A

**GW7.5 Sagittal CR 21 mm, C6202
Level 6: Slide 18, Section 2
PONS AND PERIPHERAL GANGLIA**



0.5 mm

See level 6 in Plates 212 A-D.

PLATE 220B

↑ Arrows indicate the presumed *direction of neuron migration* from neuroepithelial sources.

↑ Arrows indicate the presumed *direction of axon growth* from peripheral nerves.

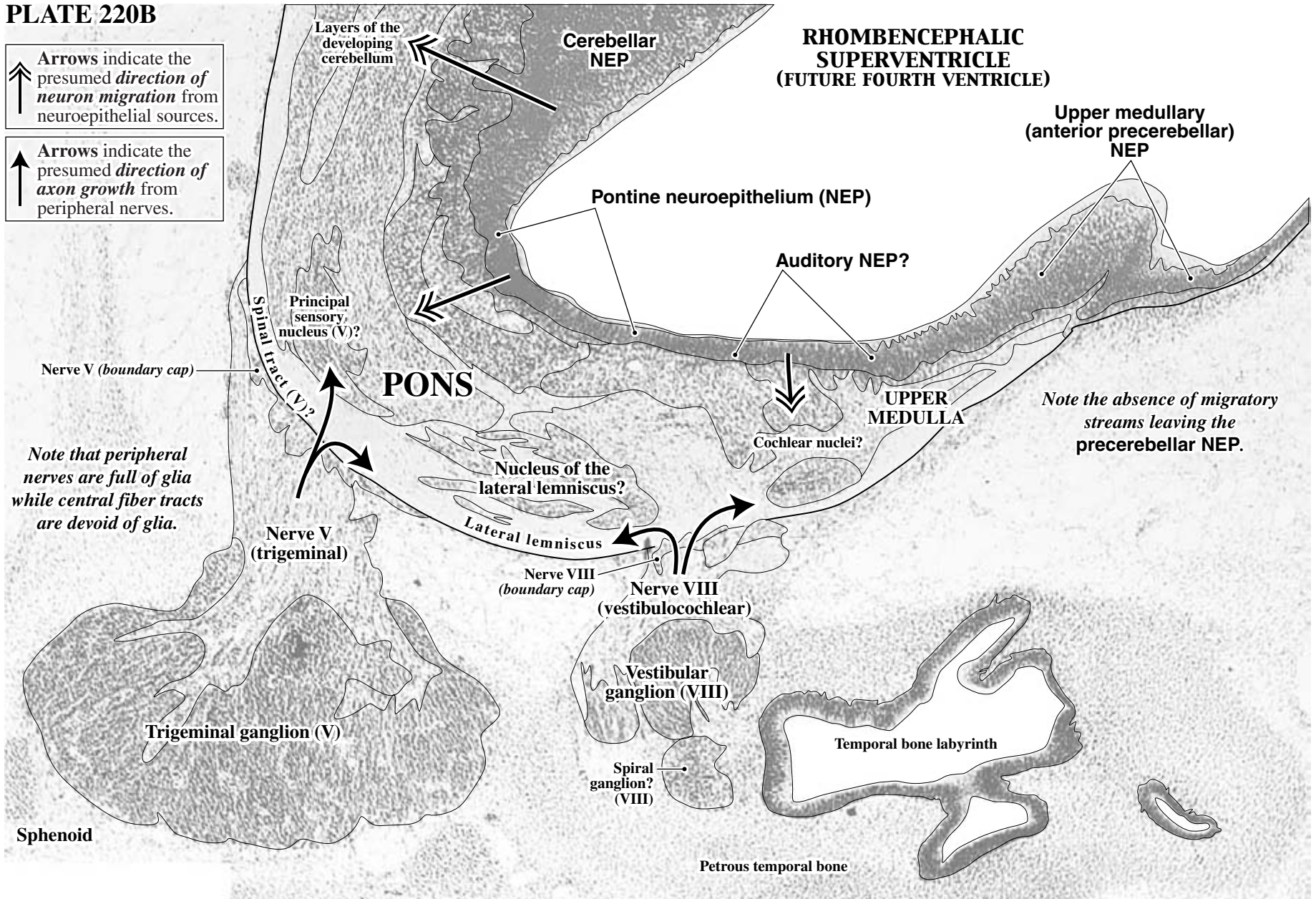
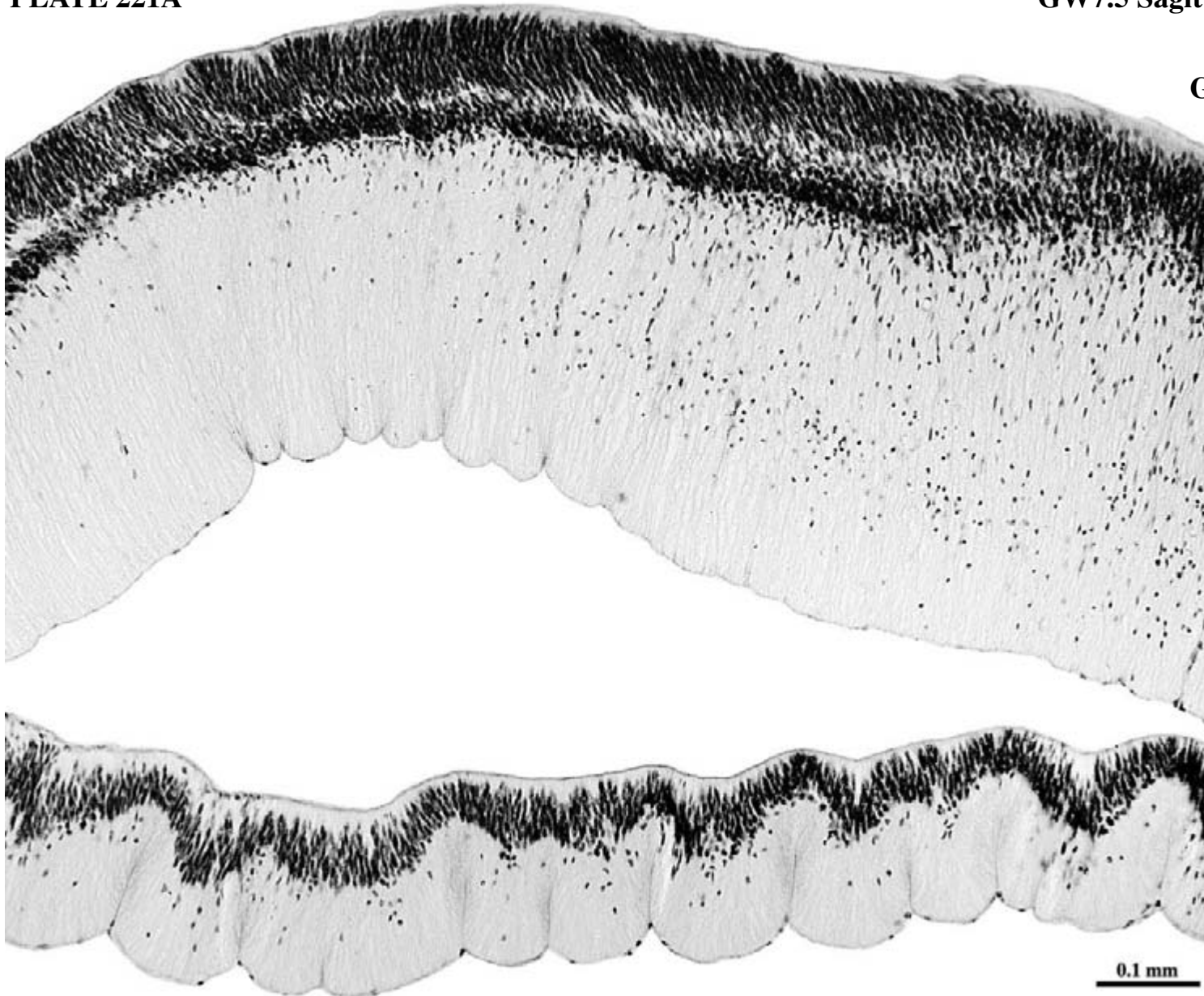


PLATE 221A

**GW7.5 Sagittal CR 21 mm, C6202
Medial to Level 1
MIDLINE RAPHE
GLIAL STRUCTURE**



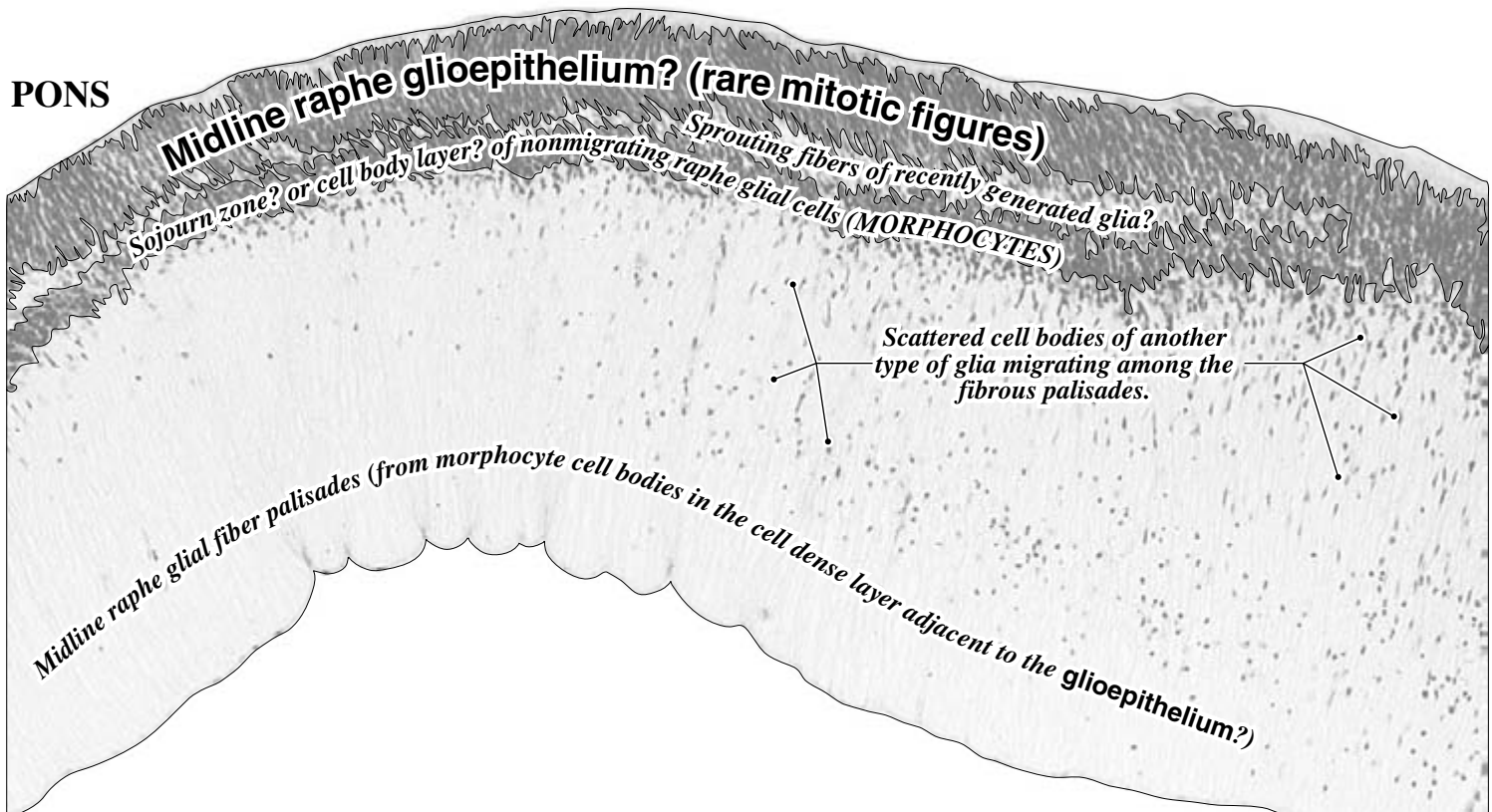
**Pons adjacent to
the pontine
flexure in slide 31,
section 2.**

**Cervical spinal
cord adjacent to
the medullary
flexure in slide 32,
section 2.**

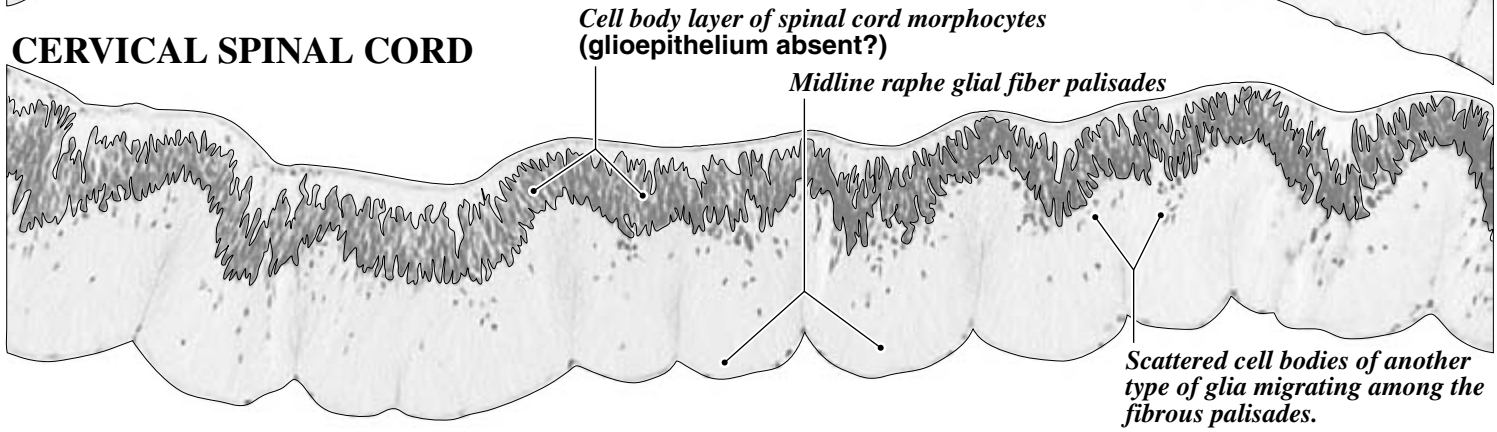
See level 1 in Plates 207A-D.

PLATE 221B

PONS



CERVICAL SPINAL CORD



The **midline raphe glial system** is prominent in regions where the shape of the brain and spinal cord sharply change curvature. Van Hartesveldt et al. (1986) described this in rats, and it is virtually identical in man. The strong fibrous palisades may provide structural stability in the region of these curvatures. Consequently, we call the specialized glia **MORPHOCYTES**.

There is structural variability between the pons and spinal cord. In the **PONS** (top panel) there are two cell-dense layers near the ventricular surface. Since mitotic figures are rare in the layer adjacent to the ventricle, it may not be an active germinal zone generating glia. However, at earlier stages of development (to be described in Volume 5), this layer is full of stem cells generating midline raphe glia. The second layer is most likely the densely packed cell bodies that have long fibrous processes extending to the pial surface rather than a premigratory sojourn zone. Morphocytes may be predominantly nonmigratory cells that differentiate at the site of their generation (similar to the ependymal cells that eventually line the ventricular system).

In the **SPINAL CORD** (bottom panel) there is one cell-dense layer adjacent to the central canal. These are the cell bodies with relatively short (compared to the pons) fibrous processes that extend to the pial surface.

In both regions, there are widely scattered cells dispersed between the fibers. These are most likely other types of glial cells that do not play a structural role (nonmorphocytes) in the raphe system.

GLOSSARY

An asterisk in front of a term indicates that it is a separate entry in the Glossary with additional information. Terms referring to transient developmental structures are underlined.

A

Abducens nucleus (VI) – An aggregate of cranial nerve motor neurons situated beneath the fourth ventricle in the *pons. The nucleus receives input from the *vestibular nuclear complex and is the source of motor fibers of cranial *nerve VI that innervate the lateral rectus muscle of the eye.

Accessory nucleus (XI) – A column of motor neurons that extends from the region of the *nucleus ambiguus in the medulla to segments 5-6 of the cervical spinal cord. Its axons form *nerve XI that innervates the sternocleidomastoid and trapezius muscles.

Accumbent NEP – Neuroepithelial source of the neurons and neuroglia of the *nucleus accumbens. After cessation of neurogenesis, this germinal matrix is transformed into a *glioepithelium.

Alisphenoid process – A cartilaginous structure in the developing skull that surrounds part of the trigeminal nerve and the facial nerve; eventually it becomes the posterior wing of the *sphenoid bone.

Allocortex (embryonic) – Portion of the cerebral hemispheres that, in contrast to the *neocortex, develops into a “three-layered” (oligolaminar) cortex. Among prominent allocortical regions are the *hippocampus, the *primary olfactory cortex, and the *entorhinal cortex.

Ammon’s horn (hippocampus) – Part of the *hippocampus that contains a prominent layer of large pyramidal cells.

Ammonic migration and sojourn zone – Migrating pyramidal cells originating in the *ammonic NEP that settle in *Ammon’s horn.

Ammonic NEP – Neuroepithelial patch of the *hippocampal NEP, the putative source of the pyramidal cells of Ammon’s horn.

Amygdala – A large subcortical structure with several subdivisions in the temporal lobe, implicated in adults in behavioral aggression.

Amygdaloid NEP – Neuroepithelial lining of the anterior wall of the posteroventral *telencephalic superventricle, the presumptive site of origin of neurons and glia of the *amygdala. It is continuous rostrally with the posterior *striatal NEP and caudally with the ventral *hippocampal NEP.

Amygdalohippocampal area – Also called the cortical amygdaloid transition area, connected in the mature brain with the *bed nucleus of the stria terminalis and the *ventromedial hypothalamic nucleus.

Ansa lenticularis – Fiber tract that originates in the internal (medial) segment of the *globus pallidus, courses dorsal to the *zona incerta, and terminates in the *thalamus, in particular the *ventral complex and the *centromedian nucleus.

Anterior amygdaloid area – A region of small to medium-sized cells in the mature *amygdala that represents a transition zone between the *substantia innominata and the amygdaloid complex proper.

Anterior commissure (embryonic) – Large fiber bundle that crosses in the ventral telencephalon and interconnects in the mature brain several forebrain structures on the right and left sides, including the *primary olfactory cortex, the *entorhinal cortex, the *amygdala, and some components of the *temporal lobe. Some of its fibers have crossed the midline by GW9.

Anterior complex (thalamus) – A group of anterior thalamic nuclei with related connections. Components of the anterior thalamic complex are the anterodorsal nucleus, the anteromedial nucleus, and the anteroventral nucleus. In the mature brain, the afferents of the anterior complex come principally from the *hippocampal region and the *mammillary body. The ascending efferents terminate in the *cingulate gyrus, while the descending efferents terminate in the *mammillary body. The anterior thalamic complex is a component of the “limbic system.”

Anterior extramural migratory stream – Large stream of young neurons migrating anteroventrally from the *anterior precerebellar NEP to the *pontine gray and the *reticular tegmental nucleus. The stream forms during the latter part of the late first trimester.

Anterior pituitary gland (embryonic) – The anterior lobe of the pituitary gland, also known as the adenohypophysis. It is derived from Rathke’s pouch during embryonic development.

Anterior precerebellar NEP – Neuroepithelial source of neurons of the *pontine gray and *reticular tegmental nucleus. It lines the *rhombencephalic supraventricle dorsally and the young neurons migrate to their target structures by way of the *anterior extramural migratory stream.

Aqueduct (embryonic) – See **Mesencephalic supraventricle**

Arcuate nucleus (hypothalamus) – A small-celled nucleus that surrounds the base of the third ventricle posteriorly. It contains releasing hormones and is involved in the central nervous regulation of the anterior pituitary gland.

Atlas – The first cervical vertebra.

Auditory germinal trigone – Proliferative site in the *rhombencephalon that contains the NEP and the external germinal layer of the *cochlear nuclei and contributes stem cells to the rhombencephalic *chordoid plexus.

Axis – The second cervical vertebra.

B

Basal ganglia – A broad term that includes three large ganglionic (subcortical) components of the telencephalon, the *caudate nucleus, the *putamen, and the *globus pallidus. The latter two are also referred to as the *striatum. Basal ganglia pathologies have been linked to Parkinsonism, Huntington’s disease, and other motor abnormalities.

Basal ganglionic NEP and SVZ – Composed of three large hillocks or eminences protruding into the *telencephalic supraventricle – the anterolateral, the antero-medial, and the posterior – and contain the neuroepithelial and subventricular stem cells that furnish neurons and neuroglia to the *basal ganglia. A fourth component is the cortico-ganglionic NEP/SVZ. The SVZ is far more prominent in the basal ganglia than in the developing *cerebral cortex.

Basal nucleus (amygdala) – The largest nucleus in the amygdala that forms a major part of the *basolateral complex. It is separated from the *lateral nucleus by a thin fibrous band.

Basal nucleus of Meynert – Large-celled component of the *substantia innominata that provides in the mature brain cholinergic input to the *cerebral cortex.

Basal occipital – The occipital bone surrounding the *foramen magnum.

Basal telencephalic NEP and SVZ – Putative source of neurons and neuroglia of the *basal nucleus of Meynert and the *substantia innominata.

Basolateral complex (amygdala) – The largest and best differentiated part of the amygdala in humans. Its principal components are the *basal nucleus and the *lateral nucleus.

Bed nucleus of the stria terminalis – A large subcortical telencephalic field with indistinct boundaries. It is situated medial to the *globus pallidus, lateral to the *septum, and is transected by the *anterior commissure; a thin portion extends back to the *amygdala adjacent to the *stria terminalis. It has its own germinal source, the *strionuclear NEP and GEP.

Boundary cap – A thin ring of proliferative cells surrounding the spinal and cranial nerves at the site they enter the central nervous system.

Brachium of the inferior colliculus – A fiber tract situated superficially in the fibrous layer covering of the *inferior colliculus. It is composed of ascending auditory fibers from the inferior colliculus and auditory nuclei in the *pons to the *medial geniculate body.

C

Calcarine sulcus (embryonic) – Cortical fissure in the occipital lobe; its wall is the target of the visual fibers from the *lateral geniculate body. It is recognizable in incipient form by the end of the first trimester.

Caudate nucleus – Elongated and arched component of the *basal ganglia beneath the *cerebral cortex. It abuts the lateral ventricle and extends from anterodorsal (its “head”) to posteroventral (its “tail”).

Cajal-Retzius cells – The earliest-generated *cerebral cortex neurons. Their perikarya are oriented parallel to the pial surface in *layer I.

Central autonomic area (spinal cord) – Region of the spinal cord that surrounds the *central canal and is implicated in nociceptive and autonomic functions. It may be continuous rostrally with the periaqueductal *central gray.

Central canal (embryonic) – Portion of the ventricular system that extends caudally from the cervical to the sacral segments of the *spinal cord. During embryonic development, the proliferative *neuroepithelium lining this canal is the source of neurons and neuroglia of the spinal cord. After the cessation of neurogenesis, the shrunken central canal is lined by the *ependyma.

Central complex (thalamus) – A group of contiguous central thalamic nuclei, including the *centromedian, central lateral, and paracentral nuclei.

Central gray (periaqueductal) – Oval shaped region in the core of the mesencephalon that surrounds the *aqueduct and is capped by the *superior colliculus and the *inferior colliculus.

Central nucleus (amygdala) – Part of the *corticomedial complex that is sometimes put in a class by itself. A large nucleus situated lateral to the medial nucleus that extends from the *anterior amygdaloid area to the caudal pole of the amygdala where it blends with the *putamen and the tail of the *caudate nucleus.

Central nucleus (inferior colliculus) – Laminated core of the *inferior colliculus where auditory fibers of the *lateral lemniscus terminate in a tonotopic order.

Centromedian nucleus (thalamus, embryonic) – Large spherical structure surrounded by fibers of the internal medullary lamina, classified with the *central complex of the thalamus. It is a paleothalamic structure that has extensive connections with the *striatum and the *reticular formation of the midbrain and is prominent by the end of the first trimester.

Cerebellar NEP – An extensive neuroepithelial matrix that lines the *rhombencephalic superventricle lateral to the *dorsal rhombic lip. It is the direct source of the neurons of the *cerebellar deep nuclei and the *Purkinje cells, and an indirect source of its basket, stellate and granule cells, by way of its secondary proliferative matrix, the *external germinal layer.

Cerebellar transitional field (CTF) – Transient cellular and fibrous layers, composed of migrating deep neurons and Purkinje cells, and of exiting and entering fiber tracts, prior to the formation of the cerebellar cortex during the second trimester.

Cerebellum (deep nuclei) – Three pairs of ganglionic structures in the depth of the cerebellum: the medial *fastigial nucleus; the intermediate *interpositus nucleus, and the lateral *dentate nucleus. The efferent fibers of cerebellar *Purkinje cells synapse with the neurons of the cerebellar deep nuclei which, in

turn, are the source of cerebellofugal fibers that terminate in structures outside the cerebellum. The early-generated deep nuclei neurons are mostly situated superficially (above the later generated Purkinje cells) during the late first trimester.

Cerebellum (hemisphere) – Portion of the cerebellum situated on either side of the midline *vermis. It is prominent in the cerebellum in higher mammals and man.

Cerebellum (vermis) – Midline portion of the cerebellar cortex. It is a relatively small part of the cerebellum in higher mammals and man.

Cerebral cortex (embryonic) – The expanding and differentiating bilateral brain tissue covering the lateral, dorsal and medial aspects of the telencephalic superventricles. It has two major components, the *neocortex and the *allocortex. The *stratified transitional field and the *cortical plate begin to form during the late first trimester.

Cerebral peduncle (embryonic) – Fibrous region along the ventrolateral aspect of the diencephalon and mesencephalon, containing fibers of the *corticofugal tract. It begins to form by the end of the first trimester and some of its fibers reach the formative *pontine gray.

Choroid plexus (embryonic) – Glycogen-rich epithelial tissue that forms and begins to expand during the later first trimester in the *rhombencephalic and *telencephalic superventricles. It is formed by proliferative stem cells associated with the cerebellar *germinal trigone and an analogous germinal site in the *hippocampus. The fetal choroid plexus may play a role in the anaerobic metabolism of the early developing brain. During late-fetal development, it becomes gradually transformed into the mature choroid plexus of the shrunken fourth ventricle, with a different cellular composition and perhaps a different function.

Cingulate cortex – A rostrocaudally extending medial region of the cerebral cortex with an *allocortical organization. By virtue of its structure and principal connections, it is considered a component of the “limbic system.”

Cingulate NEP – Long neuroepithelial stretch along the medial bank of the *telencephalic superventricle that generates the neurons and glia of the *cingulate cortex.

Clastrum (embryonic) – Subcortical gray matter adjacent to the *insula. During embryonic development, it is in the path of the *lateral migratory stream.

GLOSSARY

Cochlear NEP – Neuroepithelial patch in the vicinity of the *ventral rhombic lip, the putative source of neurons and glia of the ventral and dorsal *cochlear nuclei.

Cochlear nuclei – The dorsal and ventral auditory nuclei are targets of *nerve VIII auditory fibers. Both nuclei contribute axons to the *lateral lemniscus that terminate in the *nuclei of the lateral lemniscus, the *inferior colliculus, and the *medial geniculate body.

Cortical NEP – Extensive and continuous neuroepithelial lining of the lateral, dorsal and medial banks of the *telencephalic supraventricle. It is the sole constituent of the *cerebral cortex during the early embryonic period but, following a strict timetable and spatial gradient, it expands and then shrinks as a class of differentiating neurons and glia leave it to enter the *stratified transitional field and migrate to the *cortical plate. The cortical NEP is also the source of a secondary proliferative matrix, the *subventricular zone, and of a fate-restricted *glioepithelium and the cells that line the enduring *ependyma.

Cortical nucleus (amygdala) – Also called the periamygdaloid cortex, part of the *corticomedial complex in the superficial amygdala.

Cortical plate – The densely packed cellular band in the embryonic and fetal *cerebral cortex that later becomes the laminated *gray matter. It is situated between the future molecular layer, or *layer I and the subplate (the future layer VII).

Corticofugal fibers (embryonic) – Collective term for the efferent fiber system (the traditional pyramidal tract) that originates in the cerebral cortex and terminates in subcortical structures. It is known by different names along its path from rostral to caudal: *internal capsule, *cerebral peduncle, *transpontine corticofugal tract, and corticospinal tract. The corticofugal tract begins to form during the end of the late first trimester but its fibers do not reach the spinal cord before the end of the second trimester.

Corticoganglionic NEP – A transitional neuroepithelium at the junction between the *cortical NEP and the *basal ganglionic NEP and SVZ.

Corticomedial complex (amygdala) – Portion of the *amygdala that includes the *anterior amygdaloid area, the *central nucleus, and the *cortical nucleus.

Cricoid cartilage – A ring-shaped cartilage that forms the base of the larynx.

Cuneate fasciculus – A large fiber tract in the dorsolateral *spinal cord. It is composed of the ascending branch of the primary sensory axons of dorsal root ganglion cells that terminate topographically in the *cuneate nucleus.

Cuneate nucleus (embryonic) – The target of the *cuneate fasciculus in the lower *medulla and the source of second-order somatosensory fibers that cross the midline and enter the contralateral *medial lemniscus. It is recognizable by GW8.

D

Dentate gyrus – Curved small-celled component of the *hippocampus, interlocked with the large-celled *Ammon's horn. Although the *dentate migration is recognizable by the end of the first trimester, the blades of the dentate gyrus do not form until the second trimester.

Dentate migration – Precursors of granule cells of the *dentate gyrus that migrate from a patch of the *hippocampal NEP to form what will later become the secondary germinal matrix of the hippocampus, the *subgranular zone.

Dentate nucleus (embryonic) – Lobulated and largest of the *cerebellar deep nuclei in the core of the cerebellar hemispheres, it is the principal source of efferent fibers of the *superior cerebellar peduncle. In the embryonic cerebellum, the early-generated neurons of the dentate nucleus are situated superficially and do not descend until after the late-generated *Purkinje cells migrate toward the surface to form the cerebellar cortex toward the early fetal period.

Diagonal band of Broca – Oblique nucleus situated ventral to the medial *septum. It is subdivided into a vertical limb dorsally and a horizontal limb ventrally.

Diencephalic NEP – The extensive neuroepithelial lining of the *diencephalic supraventricle. Its different mosaic components, distinguished by their bilaterally symmetrical variable thicknesses and evaginations or invaginations into the ventricle, are the source of neurons and glia of the different nuclei of the *diencephalon.

Diencephalic supraventricle – Large midline component of the embryonic ventricular system that is later reduced to the small and narrow third ventricle. It is confluent laterally, by way of the foramen of Monro, with the *telencephalic supraventricle, and caudally with the *mesencephalic supraventricle. Its lining, the *diencephalic NEP, is the source of all the neurons and glia of the *diencephalon.

Diencephalon (embryonic) – Extensive forebrain region flanked laterally by the telencephalon and continuous caudally with the mesencephalon. Among its larger components are the *epithalamus and the *thalamus dorsally; the *preoptic area, *hypothalamus, and *subthalamus ventrally; and the *pretectum caudally. Its development precedes that of dorsal *telencephalon.

Dorsal complex (thalamus) – Collective term for two structurally and functionally related dorsally situated thalamic regions, the *dorsomedial nucleus and the dorsolateral nucleus.

Dorsal funiculus – *See Dorsal white matter*

Dorsal gray matter (spinal cord) – Wing-shaped region of the spinal gray matter, the target of the local collaterals dorsal root afferent fibers. Its principal component is the small-celled substantia gelatinosa. The neurons of the dorsal gray matter originate in the *neuroepithelium flanking the transient dorsal spinal canal.

Dorsal hippocampus (embryonic) – The portion of the *hippocampus located above the thalamus during the early stages of embryonic development. In the late second trimester, this region is displaced by the expanding *cerebral cortex laterally and then ventrally.

Dorsal motor nucleus (X) (embryonic) – An early-forming column of parasympathetic preganglionic motor neurons dorsolateral to the *hypoglossal nucleus. Their axons leave the brain in cranial *nerve X and terminate in the parasympathetic ganglia supplying the viscera of the thoracic, pericardial, and abdominal cavities. *See also Nerve X.*

Dorsal rhombic lip – The dorsolateral anchoring point of the *medullary velum that covers the expanding *rhombencephalic supraventricle. Following the formation of the cerebellar *external germinal layer by the end of the first trimester, it is recognized as the *germinal trigone. The *ventral rhombic lip forms the ventrolateral anchoring point of the medullary velum.

Dorsal sensory nucleus (X) (embryonic) – A medial nucleus in the early-forming *solitary nuclear complex of the *medulla that lies dorsolateral to the *dorsal motor nucleus (X). *See also Nerve X.*

Dorsal tegmental nucleus – Situated in the central gray dorsal to the *trochlear nucleus and extending caudally into the pons. It is targeted by fibers of the *mammillotegmental tract.

Dorsal white matter (spinal cord) – Medial fibrous component of the white matter situated between the wings of the *dorsal gray matter; also known as the dorsal column or the dorsal funiculus. It contains ascending somatosensory and proprioceptive fibers that terminate in the dorsal column nuclei of the medulla. In the upper spinal cord it has two distinguishable parts, the *gracile fasciculus medially, and the *cuneate fasciculus laterally.

Dorsomedial nucleus (hypothalamus) – Area situated above the more distinct *ventromedial nucleus of the hypothalamus. Its principal connections are with the *bed nucleus of the stria terminalis and the *septum.

Dorsomedial nucleus (thalamus) – Also known as the medial dorsal nucleus, this component of the *dorsal complex is situated between the internal medullary lamina and the periventricular gray. Its principal connections are with the *amygdala, the *hypothalamus, the *olfactory tubercle, and the *orbital gyrus. It is considered a paleothalamic component of the limbic system rather than as a neothalamic relay nucleus to the neocortex.

E

Endopiriform nucleus (embryonic) – Small telencephalic nucleus deep to the *primary olfactory cortex and ventral to the *claustrum. The *lateral migratory stream percolates through this nucleus during development.

Entorhinal cortex – Allocortical component of the *parahippocampal gyrus. It is bordered internally by the *subicular complex and is separated from the *neocortex in the maturing brain by the rhinal sulcus. It is the source of the perforant pathway to the *hippocampus.

Ependyma – Layer of cuboidal cells that line the lumen of the permanent brain *ventricles and *central canal after dissolution of the proliferative *neuroepithelium.

Epithalamic NEP – Neuroepithelial patch of the *diencephalic NEP, the putative source of neurons and glia of the *epithalamus.

Epithalamus – Collective term for the region of the dorsal diencephalon consisting of the *habenular nuclei, the stria medullaris, and the *habenulo-interpeduncular tract.

Ethmoid – A cranial bone that lies beneath the olfactory bulb and forms part of the nasal septum.

GLOSSARY

Eustachian tube – A canal between the oral cavity and the middle ear.

External capsule – Slender fiber band situated between the *claustrum and the *putamen.

External cuneate nucleus – Situated lateral to the *cuneate nucleus in the *medulla, also known as the accessory cuneate nucleus, it is the source of the cuneocerebellar tract. The nucleus relays somesthetic and proprioceptive information from anterior regions of the body to the *cerebellum.

External germinal layer (cerebellum) – Subpial, secondary germinal matrix of the cerebellar cortex, the source of its late-differentiating granule, stellate, and basket cells. It begins to form at the end of the first trimester and persists as a source of neurons over the surface of the human cerebellar cortex until the end of the second year of postnatal life.

F

Facial motor nucleus – A large aggregate of somatic motor neurons in the ventrolateral pons dorsal to the *superior olive complex. It is the source of the motor fibers of *nerve VII that innervate the facial mimetic muscles. Subdivisions of this nucleus innervate different facial muscles.

Fastigial nucleus (cerebellum) – A deep nucleus of the *cerebellum, also known as the medial cerebellar nucleus. It is the target of Purkinje cell axons that originate in the *vermis. Its axons contribute to the large efferent system that leaves the cerebellum.

Foramen magnum – The large opening in the occipital bone that surrounds the spinal cord.

Foramen of Monro (embryonic) – Large bilateral channels that connect the paired *telencephalic superventricles with the midline *diencephalic superventricle.

Forel's fields (embryonic) – Early differentiating subthalamic tegmental field (H1 and H2) traversed by fibers of the *ansa lenticularis, lenticular fasciculus, and subthalamic fasciculus.

Fornical GEP – Fate-restricted germinal extension of the hippocampal NEP, the gliopithelium that surrounds the fornix. It may be the germinal source of the oligodendrocytes of the fornix.

Fornix – An early forming fiber tract of the *hippocampus that distributes fibers in the mature brain to the *septum and the thalamic *anterior complex, and terminates in the *mammillary body.

Fourth ventricle (embryonic) – See **Rhombencephalic superventricle**

Frontal bone – Part of the skull that lies over the frontal lobes of the cerebral cortex.

Frontal lobe or cortex (embryonic) – Region of the developing *cerebral cortex that will grow into the large frontal lobe anterior to the central sulcus. The *orbital cortex is sometimes distinguished as a separate component.

Frontal NEP – Long stretch of the cortical neuroepithelium that is presumed to generate the cells of the *frontal lobe. Before settling in the *cortical plate, the migrating and sojourning neurons form a site-specific *stratified transitional field.

Frontal nasal process – A primordium in the immature skull of the future frontal and nasal bones.

G

Ganglionic narrows – Ventricular region where the *basal ganglionic eminences protrude into the lumen of the *telencephalic superventricle.

Germinal trigone (cerebellum) – Proliferative germinal matrix of the *dorsal rhombic lip with three prongs: the *cerebellar NEP, the *external germinal layer, and the stem cells of the rhombencephalic *choroid plexus.

Gliopithelium (GEP) – Fate-restricted transient germinal matrix in the developing brain, the presumed source of neuroglia (astrocytes and oligodendroglia) precursors. There are two types of gliopithelia, the *perifascicular GEP that surround fiber tracts, such as the *fornical GEP, and another that covers the surface of the brain, the *subpial granular layer. Gliopithelia are easiest to recognize without special glial markers at sites of considerable distance from neuronal aggregates or their migratory routes.

Gliopithelium/Ependyma (G/EP) – Transient proliferative linings of the ventricle that endure into adulthood and give rise to both glia and cells of the *ependyma.

Globus pallidus – Component of the *striatum, situated medial to the *putamen, with an external (lateral) and internal (medial) segment. Pallidal fibers are the principal efferents from the striatum to the *thalamus, the *subthalamus, and the *tegmentum.

Gracile fasciculus – A large fiber tract in the dorsomedial *spinal cord. It is composed of the branch of axons of dorsal root ganglia that convey somatosensory input

from lower parts of the body. The fibers terminate topographically in the *gracile nucleus.

Gracile nucleus (embryonic) – Gray mass in the core of the *gracile fasciculus that receives input from that fiber tract. Axons of these neurons cross the midline in the *medulla, enter the contralateral *medial lemniscus. It is recognizable by GW8.

Gray matter (embryonic) – General term for the laminated component of the mature cerebral cortex with a high concentration of neuronal cell bodies and nerve processes but few myelinated fibers. In the embryonic cerebral cortex, the unlaminate cortical gray matter is known as the *cortical plate.

H

Habenular nuclei (complex) – Mediodorsally situated nuclei in the posterodorsal *thalamus; sometimes distinguished from the thalamus proper as the *epithalamus. Its two distinct components are the medial and the lateral habenular nuclei. Habenular afferents come principally from the *septum and the *hypothalamus, and its efferents form the *habenulo-interpeduncular tract.

Habenulo-interpeduncular tract (embryonic) – An early forming fiber bundle, also known as the fasciculus retroflexus, that originates in the *habenular nuclei. It courses through the posterior *thalamus and terminates in the *interpeduncular nucleus of the midbrain.

Hippocampal NEP – Long medial stretch of the *cortical NEP, the putative neuroepithelial source of neurons and glia of the hippocampus. It has three distinctive parts, the *ammonic NEP, the *dentate migration, and the *fornical GEP.

Hippocampal region – An inclusive term (also called the hippocampal formation) that includes not only the *hippocampus proper but also the *subicular complex and other components of the *parahippocampal cortex.

Hippocampus – A distinctive allocortical (three-layered or oligolaminar) region formed by the interlocking *dentate gyrus and *Ammon's horn. The hippocampus is continuous with the *subicular complex. The principal afferents of the hippocampus travel in the alveolar and perforant paths; its efferents leave by way of the *fimbria that join the *fornix.

Honeycomb matrix – Strands of radially oriented fibers surrounded by migrating cells that may be present by the end of the first trimester in the *superior colliculus

and forms during the second trimester in the *stratified transitional field of the sensory areas of the developing *cerebral cortex.

Hook bundle (embryonic) – Intracerebellar decussating fiber tract that originates in the deep nuclei of the cerebellum and presumed to leave it contralaterally as the uncinata fasciculus. A prominent fibrous region identified at the base of the cerebellum in GW8-GW9 specimens may be the sprouting fibers of this tract.

Hypoglossal nucleus (XII) – A column of somatic motor neurons near the floor of the *fourth ventricle in the caudal medulla. Their axons form cranial *nerve XII that innervate the intrinsic and extrinsic muscles of the tongue.

Hyoid bone – A small bone embedded in the throat muscles that lies below the tongue and above the larynx.

Hypothalamic NEP – Stretch of the *diencephalic NEP, situated posterior to the preoptic NEP and ventral to the subthalamic NEP. It is the source of neurons and glia of the nuclei of the *hypothalamus.

Hypothalamus (embryonic) – Early differentiating large diencephalic region that surrounds the ventral (or hypothalamic) portion of *diencephalic superventricle. It is continuous anteriorly with the *preoptic area and merges caudally with the midbrain *tegmentum. The hypothalamus contains a large number of discrete nuclei, among them the *suprachiasmatic nucleus, the *supraoptic nucleus, the *paraventricular nucleus, the *arcuate nucleus, the *ventromedial nucleus, the *dorsomedial nucleus, the *lateral tuberal nucleus, and the *mammillary body.

I

Inferior cerebellar peduncle – Large fiber tract, also known as the restiform body. It contains ascending afferents to the cerebellum from the *spinal cord, the *external cuneate nucleus, the *inferior olive, and the *lateral reticular nucleus.

Inferior collicular NEP – Distinctive neuroepithelial component of the *tectal NEP surrounding the posterior pool of the mesencephalic superventricle and the source of neurons of the *inferior colliculus.

Inferior colliculus – Paired inferior hillocks of the midbrain *tectum that receive primary, secondary, and higher order auditory afferents. The output of the inferior colliculus is mainly to the *medial geniculate body in the thalamus, but some axons extend to the primary auditory cortex in the *temporal lobe.

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Inferior olive – A distinctive convoluted region in the ventrolateral *medulla, with the large principal nucleus and the smaller accessory nuclei. Its axons join the contralateral *inferior cerebellar peduncle and terminate in the *cerebellar cortex as climbing fibers.

Inferior olive (embryonic) – A compact cell aggregate in the lower *medulla, formed by neurons of the *posterior intramural migratory stream during the late first trimester. Its lamination does not begin until the end of the second trimester.

Inferior vestibular nucleus – This nucleus begins caudally in the *medulla near the *external cuneate nucleus and extends rostrally along the medial border of the *inferior cerebellar peduncle.

Infundibular recess – Small recess of the third ventricle that evaginates into the *infundibulum.

Infundibulum – Stalk extending from the ventral *hypothalamus that forms a link with the pituitary gland.

Insula (insular cortex) – Large buried neocortical region that is continuous internally with the *frontal, *parietal and *temporal lobes.

Interhemispheric fissure – Longitudinal cleft that separates the two cerebral hemispheres. The corpus callosum that traversed it in the maturing brain has not yet started to form in the first trimester.

Internal capsule (embryonic) – Massive fiber tract between the *thalamus and *striatum, composed of *thalamocortical and *corticofugal fibers. It is continuous with the incipient *cerebral peduncle caudally.

Interpeduncular nucleus – Midline mesencephalic structure above the interpeduncular fossa and between the *cerebral peduncles. It is the target of fibers of the *habenulo-interpeduncular tract.

Interpositus nucleus (cerebellum) – A deep cerebellar nucleus located between the *dentate nucleus and the *fastigial nucleus. It contains a lateral and a medial group of neurons, the emboliform near the *dentate nucleus, and the globosus near the *fastigial nucleus.

Isthmal canal – Channel that interconnects the *mesencephalic and *rhombencephalic supraventricles.

Isthmal NEP – The putative source of neurons and glia of the *isthmus.

Isthmus – Transitional brain region between the *midbrain tectum, the *pons, and the *cerebellum.

L

Lateral geniculate body – Large posteroventral thalamic region, composed in the maturing brain of the laminated dorsal lateral geniculate nucleus and its capsule of ipsi- and contralateral *optic tract fibers. Its axons project, by way of the *visual radiation, to the ipsilateral *occipital lobe.

Lateral geniculate nucleus (embryonic) – The neurons of this prominent thalamic nucleus appear to be generated dorsally in a distinct neuroepithelial locus and migrate ventrolaterally where they meet the incoming *optic tract fibers.

Lateral hypothalamic area – An ill-defined fibrous region of the *hypothalamus with scattered neurons medial to the cerebral peduncle. It is traversed by many fiber tracts, including the *medial forebrain bundle.

Lateral lemniscus – The fiber tract on the lateral surface of the *pons that contains secondary auditory fibers from the dorsal and ventral *cochlear nuclei and higher-order auditory fibers from the *superior olivary complex. The dorsal and ventral *nuclei of the lateral lemniscus are embedded within the fiber tract.

Lateral migratory stream (cortical) – Tangentially migrating neurons and glia in the developing *cerebral cortex that leave dorsal *cortical NEP and migrate laterally and ventrally to the *insula, the *temporal lobe, and other telencephalic structures that lack a nearby germinal matrix. The bulk of the lateral migratory stream follows a trajectory outlined by the receding *subventricular zone between the *basal ganglia and the lateral cortex.

Lateral nucleus (amygdala) – The most lateral nucleus in the *basolateral complex of the *amygdala. Cells from the *lateral migratory stream appear to enter the nucleus at the distinctive saw-toothed lateral edge.

Lateral olfactory tract – See **Olfactory tract**

Lateral preoptic area – An anterior continuation of the *lateral hypothalamic area in the *preoptic area.

Lateral reticular nucleus – A relatively discrete group of neurons in the caudal medulla, dorsolateral to the *inferior olive. The neurons of this precerebellar relay nucleus receive topographic exteroceptive and proprioceptive afferents from the *spinal cord and project ipsilaterally to the *cerebellum via the *inferior cerebellar peduncle.

Lateral septal nucleus – An indistinct gray mass in the lateral *septum that is closely associated with the *fornix, which provides input to these neurons.

Lateral tuberal nucleus (hypothalamus) – Two or three distinct spherical masses near the inferior surface of the *lateral hypothalamic area.

Lateral ventricles – See Telencephalic supraventricle

Lateral vestibular nucleus – Also called Deiter's nucleus. A collection of large neurons lying dorsolaterally along the wall of the *fourth ventricle. It receives primary sensory input from the vestibular ganglion via cranial *nerve VIII and its large neurons are the source of the vestibulospinal tract.

Layer I (embryonic) – Cell sparse layer beneath the pia throughout the *cerebral cortex. This is the first cortical layer to develop and contains the earliest generated cortical neurons, including the *Cajal-Retzius cells.

Locus coeruleus – Aggregate of large pigmented cells in the *pons. It is the major source of ascending and descending noradrenergic fibers that are widely distributed throughout the central nervous system.

Lusian NEP – Putative source of neurons of the so-called *subthalamic nucleus (corpus Luysi) in the *hypothalamic NEP. The intramural migration of these neurons has been traced from the region of the formative *mammillary body to the level of the subthalamus dorsolaterally.

M

Mammillary body – Distinctive region in the posteroventral hypothalamus, composed of the medial and lateral mammillary nuclei. Its principal afferents are from the *septum and *subiculum that course in the *fornix; its efferents form the *mammillothalamic and *mammillotegmental tracts.

Mammillotegmental tract – Descending fiber bundle containing *mammillary body efferents to the brain stem, including the *dorsal tegmental nucleus.

Mammillothalamic tract – Ascending fiber bundle containing efferents of the *mammillary body that terminate in the thalamic *anterior complex.

Mandible – The lower jaw bone.

Maxilla – The upper jaw bone.

Meckel's cartilage – Cartilage of the first pharyngeal arch that is associated with the formation of the mandible (lower jaw bone) and the ossicles of the middle ear.

Medial accessory olive – A small nucleus in the *inferior olive complex that contains densely packed neurons along the lateral border of the medial lemniscus. It receives proprioceptive input from the spinal cord and its efferents reach the contralateral cerebellum (mainly the vermis) by way of the *inferior cerebellar peduncle.

Medial forebrain bundle – A diffuse fiber tract that extends from the *olfactory tubercle, through the *lateral hypothalamic area, to the *substantia nigra in the midbrain *tegmentum.

Medial geniculate body – Principal thalamic relay station in the auditory pathway to the *cerebral cortex. Its afferents originate in the *trapezoid body, the *superior olivary complex, the *nuclei of the lateral lemniscus, and the *inferior colliculus. Its efferents form the auditory radiation that terminates in the *temporal lobe.

Medial lemniscus – Large fiber bundle conveying tactile and other somatosensory input to the thalamus. It originates in the *gracile and *cuneate nuclei in the *medulla, crosses to the opposite side, ascends through the *pons and *midbrain, and terminates in the *ventral posterolateral and *ventral posteromedial nuclei of the thalamus.

Medial lemniscus (decussation) – Also known as the sensory decussation, it is composed of ascending fibers of the medial lemniscus that cross to the opposite side in the medulla.

Medial longitudinal fasciculus – A dorsomedial tract in the *midbrain, *pons, and *medulla that contains ascending and descending vestibular fibers coursing beneath the *oculomotor nuclear complex, the *trochlear nucleus, the *abducens nucleus, and the *hypoglossal nucleus. It turns ventrally in the posterior medulla and extends into the ventral funiculus of the cervical spinal cord.

Medial preoptic area – A rounded mass of neurons implicated in reproductive functions.

Medial septal nucleus – An indistinct nucleus in the midline septum that is continuous with the vertical limb of the *diagonal band of Broca.

Medial vestibular nucleus – Component of the *vestibular nuclear complex situated underneath the *fourth ventricle medial to the other vestibular nuclei. Its

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neurons receive primary sensory input from the vestibular ganglion via cranial *nerve VIII and project to the cerebellum.

Median preoptic nucleus – A small nucleus in the *preoptic area that forms a narrow cap around the *anterior commissure in the midline.

Medulla (embryonic) – Region of the neuraxis, also known as the medulla oblongata, surrounding the posterior *rhombencephalic superventricle and bounded by the *pons rostrally and the *spinal cord caudally. An extremely heterogeneous region containing sensory, somatic motor, and visceral motor nuclei as well as several ascending, descending, and decussating fiber tracts.

Medullary NEP – Extensive neuroepithelial site that lines the variegated caudal bank of the *rhombencephalic superventricle. Its several subdivisions are the source of neurons and glia of the different sensory, relay and motor nuclei of the medulla.

Medullary velum – Membranous roof of the expanding *rhombencephalic superventricle. Its lateral anchor to the *rhombencephalon is known as the *dorsal rhombic lip and the *ventral rhombic lip, and its inner surface serves as a substratum for the expanding rhombencephalic choroid plexus.

Mesencephalic NEP – The extensive neuroepithelium that lines the large *mesencephalic superventricle. Its major divisions are the anterior *pretectal NEP, the dorsal *tectal NEP, the ventral *tegmental NEP, and the posterior *isthmal NEP. Subdivisions are the source, among others, of neurons and glia of the *superior colliculus and *inferior colliculus dorsally, and of the *red nucleus, the *oculomotor nuclear complex, and several tegmental nuclei ventrally.

Mesencephalic nucleus (V) – Large neurons scattered along the lateral border of the *central gray of the midbrain and pons. They are primary sensory neurons that enter the brain from the periphery early in development relaying proprioceptive information from the muscles of mastication.

Mesencephalic superventricle – Greatly inflated lumen of the embryonic *mesencephalon, situated between the *diencephalic superventricle rostrally and the *rhombencephalic superventricle caudally. The connection with the latter is by way of the *isthmal canal. It shrinks in the maturing brain into the narrow aqueduct.

Mesencephalon (embryonic) – Anterior part of the developing brainstem surrounding the *mesence-

phalic superventricle consisting of the *pretectum and *tectum dorsally, and of the *tegmentum ventrally. Among the intermediate components of the mesencephalon are the *central gray, the *oculomotor nuclear complex, the *red nucleus and the *reticular formation.

Meyer's loop – Part of the *visual radiation that takes a sharp curve in the *temporal lobe as it proceeds to the *occipital lobe.

Midbrain – *See Mesencephalon*

Middle cerebellar peduncle – Massive tract of *ponto-cerebellar fibers that originate in the *pontine gray and enter the cerebellum posterolateral to the *inferior cerebellar peduncle. The earliest fibers of this system emerge at the end of the first trimester.

Motor nucleus, V – *See Trigeminal, motor nucleus*

Motor nucleus, VII – *See Facial, motor nucleus*

N

Nasal conchae – Scroll-like processes of the *ethmoid and *maxilla that project into the lateral nasal cavity.

Neocortex (embryonic) – Portion of the cerebral hemispheres that develops a “six-layered” (multilaminar) cortical *gray matter. Neocortical development begins with the expansion of the primordial *cortical NEP devoid of differentiated neurons. It is followed by the formation of the *cortical plate, the *subventricular zone, and the different layers of the *stratified transitional field, in association with the ingrowth of *thalamocortical fibers, the outgrowth of *corticofugal fibers and the onset of the formation of intracortical connections. The principal divisions of the neocortex are the *frontal lobe, the *paracentral lobule, the *parietal lobe, the *temporal lobe, and the *occipital lobe.

Neocortical NEP – Extensive *neuroepithelium that lines the lateral and dorsal aspects the *telencephalic superventricles. The proliferating neocortical NEP cells are the source of neurons and glia that migrate to the nearby *cortical plate by way of the *stratified transitional field. Some of its cells may move to more distant sites by way of the *lateral migratory stream.

Nerve I – *See Olfactory nerve*

Nerve II – *See Optic nerve*

Nerve III (oculomotor) – Cranial motor nerve originating in the *oculomotor nuclear complex. It innervates

all the extraocular muscles – except the lateral rectus and superior oblique – and the skeletal muscles of the eyelid, the smooth sphincter muscles of the iris, and the ciliary muscles of the lens.

Nerve IV (trochlear) – Cranial motor nerve composed of axons of the *trochlear nucleus that innervates the superior oblique muscle of the eye. This nerve is unique because it exits from the dorsal surface of the *midbrain behind the *inferior colliculus.

Nerve V and ganglia (trigeminal) – A mixed sensory and motor cranial nerve that has three peripheral branches, the ophthalmic, the maxillary, and the mandibular. All three branches contain peripheral sensory fibers from the trigeminal ganglion that terminate in the *trigeminal principal sensory nucleus, the *trigeminal spinal nucleus, and the substantia gelatinosa in upper cervical segments of the *spinal cord. A bundle of fibers in the mandibular branch, originating in the *trigeminal motor nucleus, innervates the muscles of mastication.

Nerve VI (abducens) – A motor cranial nerve that originates in the *abducens nucleus and emerges near the midline at the caudal border of the *pons. The fibers innervate the lateral rectus muscle of the eye.

Nerve VII and ganglion (facial) – A mixed sensory and motor nerve, the facial nerve has three components. Primary sensory gustatory fibers from the geniculate ganglion enter the *solitary tract and nucleus. Somatic motor fibers from the *facial motor nucleus innervate the mimetic muscles. Visceral motor (parasympathetic) fibers from preganglionic neurons of the indistinct salivatory nucleus target the pterygopalatine and submandibular ganglia.

Nerve VIII and ganglia (cochlear, vestibular) – A sensory cranial nerve that contains primary auditory afferents from the spiral ganglion in the cochlea and primary vestibular afferents from the vestibular (Scarpa's) ganglion. The auditory afferents terminate in the dorsal and ventral *cochlear nuclei; the vestibular afferents terminate in the nuclei of the *vestibular nuclear complex and some reach the *cerebellum.

Nerve IX and ganglia (glossopharyngeal) – A mixed sensory and motor cranial nerve. The sensory part of nerve IX originates in the superior and inferior ganglia, and relays gustatory input from the posterior third of the tongue and visceral sensory input from the tonsils, the Eustachian tube, and the carotid sinus. These fibers enter the solitary tract and terminate in the *solitary nucleus. The somatic motor part of nerve IX originates in the *nucleus ambiguus and innervates the pharyngeal and laryngeal muscles. The

visceral motor fibers from parasympathetic preganglionic neurons in the salivatory nucleus terminate in the otic ganglion.

Nerve X and ganglia (vagus) – A mixed sensory and motor nerve, with some somatic and many visceral afferents and efferents associated with the cranio-sacral parasympathetic ganglia. The sensory fibers originate peripherally in the superior and inferior ganglia and are widely distributed throughout the body, including the pharynx, larynx, trachea, esophagus, and all the thoracic and abdominal viscera. They terminate centrally in the *solitary nucleus and at other medullary sites. Most of its preganglionic motor neurons are located in the *dorsal motor nucleus (X).

Nerve XI (accessory) – This motor nerve has a cranial and a spinal component. The cranial fibers originate in the *nucleus ambiguus and innervate the muscles of the larynx and pharynx. The spinal motor fibers originate in a motor column of the cervical *spinal cord and innervate the sternocleidomastoid and upper trapezius muscles.

Nerve XII (hypoglossal) – A somatic motor cranial nerve that originates in the *hypoglossal nucleus and innervates the intrinsic and extrinsic muscles of the tongue.

Neuroepithelium (NEP) – Pluripotential pseudostratified tissue of neural stem cells that extends from the frontal pole rostrally to the last segment of the spinal cord caudally and is the source of all neurons and neuroglia of the developing central nervous system. The NEP cells initially form the neural plate, then fold dorsally and fuse to form the neural tube (future spinal cord) caudally and the variegated cephalic vesicles (the future brain) rostrally. After that closure, the lumen of the cephalic vesicles expands enormously to form the *rhombencephalic, *mesencephalic, *diencephalic, and *telencephalic superventricles. This expansion provides the space for the mitotic division of NEP cell nuclei that have to shuttle to the fluid-filled lumen to undergo mitosis. The continuous but variegated cephalic NEP lining the ventricles has a mosaic organization, being composed of bilaterally symmetrical long and intermediate stretches, and short patches that give rise to neurons and glia of different brain regions, distinct structures, and specific cell types. Examples of long stretches are the *cortical NEP and the *cerebellar NEP; of intermediate stretches, the *thalamic NEP and the *hypothalamic NEP of the inclusive *diencephalic NEP; and of short patches, the *ammonic NEP and *dentate NEP of the inclusive *hippocampal NEP. The primary NEP is also the source of several *secondary germinal matrices that generate microneurons with locally arboriz-

ing axons. Finally, as neurogenesis winds down the pluripotential NEP is transformed at many sites into a *glioneithelium, such as the *fornical GEP, or into the *ependyma that lines the enduring ventricles.

Nuclei of the lateral lemniscus – Both components of this system, the dorsal nucleus and the ventral nucleus, receive their major input from the *cochlear nuclei and the *superior olivary complex by way of the *lateral lemniscus. Both nuclei are connected with the *inferior colliculus and the *medial geniculate body.

Nucleus accumbens – Ganglionic component of the ventral telencephalon ventromedial to the *striatum. It is distinguished from the striatum by its cellular organization, molecular composition, and intimate connections with the *hypothalamus, *amygdala, and other regions of the limbic system.

Nucleus ambiguus – Aggregate of somatic motor neurons that form a thin column in the ventrolateral medulla. Its axons innervate the muscles of the larynx and pharynx via *nerve IX.

O

Occipital lobe or cortex (embryonic) – Posterior region of the developing cerebral cortex that will be the target of *visual radiation fibers from the *lateral geniculate body and the *pulvinar.

Occipital NEP – Putative neuroepithelial source of the neurons and glia of the occipital lobe. It is flanked in the fetal neocortex by the occipital *subventricular zone and the *stratified transitional field.

Oculomotor nerve – *See Nerve III*

Oculomotor nuclear complex – Situated at the base of the periaqueductal *central gray, the cell columns of this complex extend from the anterior pole of the *superior colliculus rostrally to the *trochlear nucleus caudally. Its somatic motor nuclei innervate the medial rectus, inferior rectus, superior rectus and inferior oblique muscles of the eye, and are associated with the fibers of the *medial longitudinal fasciculus. Most prominent of its autonomic (preganglionic) components is the dorsally located Edinger-Westphal nucleus.

Odontoid process – A tooth-like projection on the superior surface of the *axis that articulates with the *atlas.

Olfactory bulb – Laminated brain structure where the first-order fibers of the *olfactory nerve terminate and the second-order fibers of the *olfactory tract orig-

inate. It is composed of three classes of neurons: large mitral cells, the intermediate tufted cells, and the small granule cells.

Olfactory NEP – Putative source of the earlier-generated mitral and tufted cells of the olfactory bulb that lines the olfactory recess of the *telencephalic superventricle. The later-generated granule cells are supplied by the *rostral migratory stream.

Olfactory nerve (embryonic) – Composed of the fine axons of bipolar cells in the olfactory epithelium that terminate in the *olfactory bulb. The nerve is recognizable in GW7.5 specimens.

Olfactory tract – Large fiber bundle of second-order fibers that originate in the *olfactory bulb with two parts, the larger lateral olfactory stria and the smaller medial stria. The fibers of the lateral stria terminate in the *olfactory tubercle, the *primary olfactory cortex, and *corticomedial complex of the amygdala.

Olfactory tubercle – Allocortical area in the ventral telencephalon between the *diagonal band of Broca and the *nucleus accumbens. Its input comes mainly from the lateral *olfactory tract.

Optic chiasm (embryonic) – Site of crossing of fibers of the *optic nerve. Fibers from the nasal half of each retina cross here to the opposite side while those from temporal half proceed uncrossed. The earliest crossing fibers appear in GW7.5 specimens.

Optic nerve – This large fiber tract contains the axons of retinal ganglion cells. Beyond the *optic chiasm it is called the *optic tract.

Optic tract – Large bundle of crossed and uncrossed retinal afferent fibers. In the human brain the majority of the fibers terminate in the *lateral geniculate body; others proceed to the *superior colliculus, the *suprachiasmatic nucleus, the *pretectum, and some other structures.

Orbital cortex – Ventromedial region of the *frontal lobe with afferents from the thalamic dorsomedial nucleus and efferents to the *preoptic area and the *hypothalamus.

Orbitofrontal NEP – Putative source of neurons and glia of the orbitofrontal cortex.

Orbitosphenoid – A cartilaginous structure in the developing skull that surrounds the *optic nerve and eventually becomes the lesser wing of the sphenoid bone.

P

Palatal process – Part of the *maxilla that forms the hard palate in the roof of the mouth.

Pallidum – *See Globus pallidus*

Parabrachial nucleus – Dorsolateral pontine structure with indistinct boundaries that surrounds the *superior cerebellar peduncle. Its principal input comes from the *solitary nucleus and its efferents target the *ventral posteromedial nucleus of the thalamus, the *amygdala, and the *insular cortex.

Paracentral lobule (embryonic) – Incipient cortical region that becomes later divided by the central sulcus into the precentral gyrus and the postcentral gyrus. The term is used to distinguish this site of the presumptive motor and sensory projection areas from the *frontal lobe anteriorly and the *parietal lobe posteriorly.

Paracentral NEP – Putative neuroepithelium of the *paracentral lobule in the developing neocortex. It is flanked by the paracentral *subventricular zone and the paracentral *stratified transitional field.

Parahippocampal cortex (embryonic) – Allocortical and neocortical region between the *hippocampus and the *temporal lobe that will become the parahippocampal gyrus. Its subdivisions are the *subicular complex and the *entorhinal cortex.

Parahippocampal NEP – Putative source of the neurons and glia of the parahippocampal gyrus. It is flanked by the parahippocampal *subventricular zone and the parahippocampal *stratified transitional field.

Paraventricular nucleus (hypothalamus) – Prominent neuroendocrine structure abutting the third ventricle with a magnocellular and a parvocellular division. The large neurons of the paraventricular nucleus are the source of oxytocin and vasopressin that reach the posterior pituitary gland by axoplasmic flow. The small neurons of the nucleus are the source of releasing hormones conveyed to the portal vessels of the median eminence.

Parietal bone – A cranial bone in the skull that eventually lies over the parietal lobe of the cerebral cortex.

Parietal lobe of cortex (embryonic) – Region of the developing neocortex bounded anteriorly by the *paracentral lobule and posteriorly by the *occipital lobe.

Parietal NEP – Long stretch of the cortical neuroepithelium containing the neural stem cells of the *parietal

lobe. It is flanked by the parietal *subventricular zone and *stratified transitional field.

Perifascicular GEP – Fate-restricted gliopithelium, the presumed source of oligodendrocytes that surround a fiber tract, such as the *fornical GEP.

Periventricular complex (thalamus) – The thalamic region surrounding the *mesencephalic superventricle that will become partitioned as the paracentral, parafascicular, paratenial, paraventricular, and reuniens nuclei. Its principal connections are with limbic system; connections with the *neocortex are sparse.

Petrous temporal bone – Part of the temporal bone that contains the internal ear and semicircular canals.

Pineal gland – Midline endocrine gland connected by its stalk to the pineal recess of the dorsal *mesencephalic superventricle. It secretes melatonin and other indoleamines. It is believed to receive indirect visual input from the retina.

Piriform cortex – *See Primary olfactory cortex*

Pituitary gland – *See Anterior pituitary gland; Posterior pituitary gland*

Pons (embryonic) – Developing brainstem region, situated between the *isthmus and the *medulla, that surrounds the anterior part of the *rhombencephalic superventricle. It contains some early ascending, descending and decussating fiber tracts, the sensory and motor nuclei of some of the cranial nerves, and the *reticular formation.

Pontine gray (embryonic) – This massive basal region of the *pons is just beginning to form as neurons of the *anterior extramural migratory stream start to settle and the earliest descending *corticofugal fibers reach the site. Corticofugal axons that collateralize here are the principal afferents of the pontine gray neurons that are, in turn, the source of the pontocerebellar fibers of the *middle cerebellar peduncle.

Posterior commissure (embryonic) – Early forming decussating fiber tract in the dorsal *mesencephalon that interconnects several nuclei in the *pretectum and *tectum.

Posterior complex (thalamus) – Division of the thalamus that includes the *lateral geniculate body, the *pulvinar, and the *medial geniculate body. The neurons of the lateral geniculate body and pulvinar appear to originate in a distant source and migrate from dorsal to ventral over an extended period.

GLOSSARY

Posterior intramural migratory stream – Stream of young neurons that migrate inside the parenchyma from their source in the posterior *precerebellar NEP dorsally, to form the *inferior olive in the ventral medulla.

Posterior extramural migratory stream – Stream of young neurons that originate in the posterior *precerebellar NEP, migrate outside the parenchyma, cross the midline ventrally, and settle on the opposite side to form the contralateral *external cuneate nucleus and *lateral reticular nucleus.

Posterior pituitary gland – The posterior lobe of the pituitary gland, also known as the neurohypophysis, is the terminal and storage site of hypothalamic neurosecretory cells.

Precerebellar NEP – Dorsally situated neuroepithelium that lines the *rhombencephalic supraventricle in the vicinity of the ventral rhombic lip and is the source of neurons of the *precerebellar nuclei. Neurons of its rostral division migrate in the *anterior extramural migratory stream and settle in the *pontine gray and the *reticular tegmental nucleus. Neurons of its posterior division form two migratory streams, the *posterior intramural migratory stream that forms the *inferior olive, and the *posterior extramural migratory stream that crosses to the opposite side and forms the *lateral reticular nucleus and the *external cuneate nucleus.

Precerebellar nuclei – A series of nuclei in the *medulla and *pons that provide massive higher-order input to the *cerebellum, including the *inferior olive, the *external cuneate nucleus, the *lateral reticular nucleus, and the *pontine gray.

Premammillary area – Region with ill-defined boundaries anterior to the *mammillary body in the *hypothalamus.

Preoptic area (embryonic) – Early developing midline region surrounding the preoptic recess of the *diencephalic supraventricle. It is contiguous anteriorly with the ventral telencephalon and blends posteriorly with the anterior *hypothalamus. It is implicated in the regulation of sexual behavior and reproductive functions.

Prepositus nucleus – Situated in the dorsomedial *medulla, it extends from the anterior part of the *hypoglossal nucleus to the posterior part of the *abducens nucleus.

Presubiculum – *Allocortical component of the *parahippocampal gyrus between the *subiculum and the *parasubiculum.

Pretectum – Dorsal area between the posterior *thalamus and the *superior colliculus with an early forming fiber system, the *posterior commissure.

Primary olfactory cortex – Allocortical region, also called the piriform lobe, where fibers of the lateral olfactory tract terminate. It is situated rostral to the *entorhinal cortex and includes the prepiriform area along the rhinal fissure and the periamygdaloid area.

Primordial plexiform layer (cortical) – The first transitional layer to appear outside of the *cerebral cortical NEP that contains *Cajal-Retzius cells and *subplate neurons. The *cortical plate forms within its boundaries later on in development.

Principal sensory nucleus (V) – *See Trigeminal, principal sensory nucleus*

Pulvinar (thalamus) – Large nucleus of the thalamic *posterior complex. Its subdivisions send fibers to various regions of the *parietal lobe, *occipital lobe, *temporal lobe, and frontal lobe. It has been implicated in multisensory integration.

Purkinje cells (embryonic) – These neurons, which form a monolayer in the maturing cerebellar cortex, are generated in the *cerebellar NEP towards the end of the first trimester, subsequent to the production of the *cerebellar deep nuclei neurons. Hence they are initially situated beneath the layer of deep neurons adjacent to the cerebellar NEP. Later they migrate toward the surface of the formative cerebellar cortex as the *external germinal layer forms there.

Putamen – Lateral component of the *striatum. It lies between the *external capsule and the *globus pallidus. It is the major source of striatal efferents to the *thalamus, *subthalamic nucleus, *substantia nigra, and *tegmentum.

R

Raphe migration – Streams of cells that originate in the dorsal *medullary NEP and are later distributed in the midline of the ventral medulla.

Raphe nuclear complex – Several smaller and some larger cell aggregates that extend in and near the midline from the *midbrain rostrally to the *medulla caudally. The raphe cells are the principal source of serotonin-containing fibers distributed along the entire neuraxis from the forebrain to the *spinal cord. They

are involved, as neuromodulators, in the regulation of sleep, wakefulness and emotional arousal.

Red nucleus (embryonic) – A prominent nucleus in the maturing brain with a small-celled (parvocellular) and a large-celled (magnocellular) division. It is recognizable during the first trimester in the vicinity of a germinal patch, identified as the *rubral NEP. The associated fibers may be early elements of the *superior cerebellar peduncle.

Reticular belt (thalamus) – Distinctive component of the thalamus; it is coextensive with the thalamic *reticular nucleus.

Reticular formation – A large collection of scattered neurons, enmeshed in a complex network of fibers, in the core of the *medulla, *pons, and *mesencephalon.

Reticular tegmental nucleus (embryonic) – Situated dorsal to the *pontine gray, this *precerebellar nucleus, also known as the nucleus reticularis tegmenti pontis, begins to form toward the end of the first trimester ahead of the pontine gray.

Reticular nucleus (thalamus, embryonic) – An early forming thin belt of cells and fibers between the wall of the *thalamus and the *internal capsule. Virtually all fibers that interconnect the thalamus and the cerebral cortex traverse the thalamic reticular nucleus.

Rhombencephalic superventricle – The greatly inflated NEP-lined lumen of the *rhombencephalon, situated between the *isthmal canal of the *mesencephalic superventricle rostrally and the central canal of the *spinal cord caudally.

Rhombencephalon (embryonic) – An extremely heterogeneous hindbrain region lining the *rhombencephalic superventricle, that includes the developing *cerebellum, *pons, and *medulla.

Rostral migratory stream – A large stream of mitotic and postmitotic cells in the forebrain extending from the cerebral *subventricular zone to the *olfactory bulb. It is a source of late generated neurons and persists after the NEP has receded or disappeared.

Rubral NEP – A distinctive neuroepithelial patch lining the *mesencephalic superventricle and situated between the *tectal NEP and the *tegmental NEP. It is the putative source of neurons of the early generated neurons of the *red nucleus.

S

Secondary germinal matrix – Layer or field of proliferative precursors of neurons and glia abutting or some distance from the primary *neuroepithelium. These fate-restricted stem cells are progeny of the NEP and persist for varying periods postnatally (some into adulthood). Examples of secondary germinal matrices are the *external germinal layer of the cerebellum, the *subventricular zone of the neocortex, the *subgranular zone of the dentate gyrus, and the *striatal subventricular zone. Typically, the secondary germinal matrices are the source of late-generated short-axoned interneurons, or microneurons.

Sella turcica – Part of the *sphenoid bone that surrounds the pituitary gland.

Septum – Midline telencephalic structure with two components, the *medial and *lateral nuclei. Its principal connections are with the *hippocampus and the *hypothalamus by way of the *fornix. The septum is a focal component of the limbic system.

Sojourn zone – Transient cellular layers formed by neurons that halt their migration for varying periods before they proceed to their final destination. They have been recognized at various sites, among them the *stratified transitional field of the cerebral cortex and the *cerebellar transitional field. It is hypothesized that the sojourn zones are sites where transient or enduring connections are made as the coarse circuitry of a brain region is established.

Solitary tract and nucleus (embryonic) – The solitary tract is an early forming medullary fiber system that may contain the primary sensory fibers of cranial *nerves VII, IX, and X that convey gustatory (VII and IX) and visceral-sensory information (IX and X) to the solitary nucleus. The *dorsal sensory nucleus of nerve X and the *commissural nucleus of nerve X are part of this nuclear complex.

Sphenoid – The skull bone that lies mainly beneath the hypothalamus, basal telencephalon, and mesencephalon. It contains the *alisphenoid and *orbitosphenoid processes and the *sella turcica around the *pituitary gland.

Spinal cord – Caudal tubular component of the central nervous system that surrounds the *central canal. Its core of gray matter (the dorsal horn, intermediate gray and ventral horn) and surrounding white matter (the dorsal, lateral and ventral funiculi) blend rostrally with the lower medulla.

Spinal nucleus (V) – *See Trigeminal, spinal nucleus*

Spinal tract (V) – *See* **Trigeminal, spinal tract**

Squamous temporal bone – The flat part of the *temporal bone that covers the *temporal lobe of the cerebral cortex.

Stratified transitional field (STF) – Transient component of the fetal *cerebral cortex, sandwiched between the *neuroepithelium and the *cortical plate. By the second trimester (GW13), illustrated in Volume 3, it has six layers of alternating cells and fibers (STF1 to STF6) that vary in their configuration in different lobes of the cerebral cortex. In the oldest specimens illustrated in this Volume (GW11), the STF has only two layers (STF1, 4/5) or at most three layers (STF1, 4, 5). The cell-rich STF4/5 is composed of sojourning neurons, and the cell-sparse STF1 of incoming *thalamocortical fibers with possibly some outgoing *corticofugal fibers.

Stria medullaris (thalamus) – Mediodorsal fiber bundle in the diencephalon coursing in an anteroposterior direction and terminating in the *habenular nuclei.

Stria terminalis – Arched fiber bundle that originates in the *amygdala, courses along the medial surface of the *caudate nucleus, and terminates in the *bed nucleus of the stria terminalis, the anterior *hypothalamus, and the *preoptic area.

Striatal NEP – Primary germinal source of neurons of the *caudate nucleus, *putamen, and *globus pallidus. It has a large anterolateral and anteromedial division, also known as the lateral and medial eminences, and a small posterior division that generates the neurons of the tail of the caudate nucleus. The posterior striatal NEP is continuous with the *amygdaloid NEP

Striatal subventricular zone (SVZ) – A massive *secondary germinal matrix flanking the striatal NEP. It generates the bulk of the neurons of *caudate nucleus, *putamen and *globus pallidus. It may also be the source of some cortical neurons.

Striatum – Term used for two components of the *basal ganglia: the *caudate nucleus and the *putamen.

Strionuclear GEP – Fate-restricted glioepithelium, the putative source of the glia of the stria terminalis, stria medullaris, and possibly other nearby fiber tracts.

Strionuclear NEP – Putative neuroepithelial source of the neurons and glia of the *bed nucleus of the stria terminalis. It is situated beneath the *striatal NEP in a notch of the *lateral ventricle near the *foramen of Monro.

Subcommissural organ – A highly vascularized circumventricular neuroendocrine organ located beneath the *posterior commissure in the roof of the *mesencephalon.

Subgranular zone (hippocampus) – A long-persisting secondary germinal matrix beneath the *granular layer of the dentate gyrus, the source of late generated dentate granule cells. It is recognizable in incipient form by the end of the first trimester.

Subicular complex – Collective term for the *parasubiculum, the *presubiculum, and the *subiculum in the *parahippocampal cortex.

Subpial granular layer – Transient cellular layer between *layer I and the pia in some regions of the developing *cerebral cortex. It may be a source of cortical astrocytes.

Subplate – A poorly-defined layer beneath the *cortical plate. It contains neurons that are the transient pioneer residents of the cortical plate.

Substantia innominata – Extensive telencephalic area with indistinct boundaries beneath the *globus pallidus. A prominent component of the substantia innominata is the *basal nucleus of Meynert.

Substantia nigra (embryonic) – A pigmented *midbrain tegmental structure in the maturing brain at the base of the *cerebral peduncle. It has two components, the dopaminergic pars compacta, and the GABAergic pars reticulata.

Subthalamic NEP – Neuroepithelial patch between the *thalamic NEP and the *hypothalamic NEP. It is also identified as the Forelian NEP to distinguish it from the *Luysian NEP.

Subthalamic nucleus – Biconvex diencephalic structure, also known as corpus Luysii, situated above the substantia nigra between the *zona incerta and the base of the *internal capsule. It has extensive reciprocal connections with the *globus pallidus, hence it is considered a component of the *basal ganglia circuitry. Subthalamic lesions produce persistent choroid movements (hemiballism) in the arms, legs and face.

Subthalamus (embryonic) – Diencephalic region situated between the *thalamus dorsally and the *hypothalamus ventrally. Its major components, *Forel's fields and the *zona incerta are recognizable in late first trimester fetuses.

Subthalamic nucleus NEP – *See* **Luysian NEP**

Subventricular zone (SVZ) – Secondary germinal matrix, derived from the primary *neuroepithelium. The SVZ flanks the NEP during early development and then abuts the ependyma when the NEP dissolves. The nuclei of proliferative SVZ cells, unlike the nuclei of NEP cells, do not shuttle to the lumen of the ventricle during mitosis. Prominent SVZs in the telencephalon are found in the *cerebral cortex and the *striatum.

Superior cerebellar peduncle (embryonic) – A large fiber tract that originates mainly in the *dentate nucleus and *interpositus nucleus. It is present by the end of the first trimester in the formative cerebellum and appears to reach the level of the *red nucleus.

Superior colliculus (embryonic) – Anterior component of the *tectum (known in lower vertebrates as the optic lobe) is a direct target of a small complement of optic nerve fibers. Several waves of migrating cells suggest its imminent lamination by the end of the first trimester. There are indications that the entering optic fibers form a *honeycomb matrix superficially.

Superior olivary complex – A group of neurons in the ventrolateral posterior *pons that receive auditory input from the dorsal and ventral *cochlear nuclei. Ipsilateral and contralateral fibers of the complex join the *lateral lemniscus and terminate in the *inferior colliculus and in the *medial geniculate body.

Superventricles – The small ventricles of the mature central nervous system – known as the lateral and third ventricles in the forebrain, the aqueduct in the midbrain, the fourth ventricle in the pons and medulla, and the central canal in the spinal cord – are a continuous system of narrow channels lined by an enduring *ependyma and filled with cerebrospinal fluid. In contrast, the ventricles of the embryonic and early fetal brain are greatly inflated, balloon-like cisterns, hence the term superventricles, lined by a continuously changing germinal matrix, the *neuroepithelium and a fetal *choroid plexus that differs from its mature counterpart. Its large and variegated components are distinguished as the *telencephalic, the *diencephalic, the *mesencephalic, and the *rhombencephalic superventricles.

Suprachiasmatic nucleus (hypothalamus) – Small, paired midline structure above the *optic chiasm. It is implicated in the photic entrainment of the circadian rhythm.

Supramammillary area (hypothalamus) – Hypothalamic region that caps the *mammillary body. Experimental studies in animals indicate that its cells project to the *dentate gyrus of the hippocampus.

Supraoptic nucleus (hypothalamus) – Located above the optic tract lateral to the optic chiasm. The large secretory neurons of this nucleus produce arginine vasopressin and oxytocin that are conveyed by axoplasmic flow to the posterior lobe of the pituitary gland.

T

Tectal NEP – Extensive mesencephalic, smooth-surfaced NEP that lines the dorsal bank of the *mesencephalic superventricle. Its larger anterior part generates the neurons and glia of the *superior colliculus, its smaller posterior part that of the *inferior colliculus.

Tectum (embryonic) – Dorsal region of the *mesencephalon, consisting of the earlier developing *superior colliculus anteriorly and the later developing *inferior colliculus posteriorly.

Tegmental NEP – The variegated ventral stretch of the *mesencephalic NEP that contains shorter patches that produce neurons and glia for various tegmental nuclei, such as the *substantia nigra and the *ventral tegmental area.

Tegmentum (embryonic) – Ventral and ventrolateral region of the *mesencephalon and *rhombencephalon with indistinct boundaries. In addition to several brainstem nuclei, it contains many early-forming ascending, decussating, and descending fiber tracts. Some tegmental nuclei have been implicated in somatomotor and visceromotor functions. The onset of development of some components of the tegmentum appear to antedate that of the *tectum.

Telencephalic superventricle – The largest component of the superventricles that begins to expand during the early first trimester and shrinks during the third trimester. It is bounded laterally, dorsally and dorsomedially by the long stretch of the *cortical NEP, and ventromedially and ventrally by the shorter *olfactory, *septal, *striatal, *hippocampal and *amygdaloid NEPs. A large portion of its lumen is occupied by the fetal telencephalic *choroid plexus.

Telencephalon (embryonic) – Extensive forebrain region consisting of both cortical and subcortical components whose neurons and glia are produced by NEP stretches and patches lining the *telencephalic superventricle.

Temporal bone – A cranial bone in the skull that covers the *temporal lobe of the cerebral cortex and contains the internal ear and semicircular canals.

Temporal lobe or cortex (embryonic) – Lateral and ventral portion of the developing cerebral cortex that will later become separated from much of the rest of the cerebral hemisphere by the lateral fissure.

Temporal NEP – Putative source of neurons and glia of the *temporal lobe. It is flanked during fetal development by the temporal *subventricular zone and the temporal *stratified transitional field.

Tenia tecta – Components of the *cerebral cortex that extend into the dorsomedial *septum (dorsal tenia tecta) and medial olfactory peduncle (ventral tenia tecta).

Thalamic NEP – Stretch of the *diencephalic NEP between the *epithalamic and the *hypothalamic NEPs. Its mosaic divisions are the putative source of neurons and glia of the diverse nuclei of the *thalamus.

Thalamocortical fibers (embryonic) – Collective term for the large afferent tracts that proceed from relay nuclei in the thalamus, by way of the *internal capsule, to the *cerebral cortex. These nuclei include the *ventral posterolateral and *ventral posteromedial nuclei, and the *lateral geniculate and *medial geniculate bodies.

Thalamus – Massive dorsal diencephalic structure with several distinct and some indistinct nuclei. As a convenience, the thalamus is divided into the following nuclear regions: the *anterior complex, the *central complex, the *dorsal complex, the *periventricular complex, the *posterior complex, the *ventral complex, and the *reticular belt.

Third ventricle – See **Diencephalic superventricle**.

Thyroid cartilage – A shield-shaped cartilage in the larynx.

Transpontine corticofugal tract (embryonic) – Portion of the large descending fiber tract in the maturing brain that traverses the *pontine gray and gives off collaterals there. Pioneering fibers of this tract are present by the end of the first trimester.

Trapezoid body – A fiber tract extending from the ventral *cochlear nucleus to the contralateral *superior olivary complex. It contains second- and higher-order auditory fibers.

Trigeminal, motor nucleus (embryonic) – Aggregate of trigeminal somatic motor neurons situated medial to the *trigeminal principal sensory nucleus. It is recognizable in late first trimester embryos.

Trigeminal, principal sensory nucleus (embryonic) – The second-order sensory neurons in the trigeminal system located dorsal and lateral to the incoming sensory root of cranial *nerve V. It receives topographic somatosensory input from the face and mouth, and its efferents cross the midline in the pons and proceed to the thalamic *ventral complex in close association with the *medial lemniscus. The nucleus is prominent by the late first trimester.

Trigeminal, spinal nucleus (embryonic) – A continuation of the *trigeminal principal sensory nucleus that extends caudally through the *medulla to the second cervical level of the *spinal cord. It is prominent by the late first trimester.

Trigeminal, spinal tract – Primary sensory fibers of the trigeminal ganglion that convey touch and pressure information from the face. The axons enter the brain in the pons and proceed caudally, forming a lateral cap around the *trigeminal spinal nucleus where they terminate in a topographic order.

Trochlear nucleus – Aggregate of somatic motor neurons located posterior to the *oculomotor nuclear complex that innervate the superior oblique muscle of the eye by way of cranial *nerve IV.

V

Ventral anterior nucleus (thalamus, embryonic) – The ventral anterior nucleus, to be distinguished from the *anteroventral nucleus (which is part of the *anterior complex of the thalamus) is the most rostral component of the thalamic *ventral complex. Its afferents come mostly from the *globus pallidus and the *substantia nigra, and its efferents terminate in the *paracentral lobule of the neocortex. The nucleus is recognizable during the late first trimester as a developing structure with putative migrating and sojourning neurons and sprouting fiber bundles.

Ventral complex (thalamus, embryonic) – A group of structurally and functionally related ventrolateral and ventral nuclei of the thalamus, including the *ventral anterior, *ventral lateral, *ventral posterolateral, and *ventral posteromedial nuclei. The ventral thalamic complex is the principal topographically organized relay system of the direct (lemniscal) and indirect (cerebellar and striatal) somatosensory and proprioceptive input system to the sensory and motor areas of the *neocortex. Its components are recognizable during the late first trimester as formative structures with putative migrating and sojourning neurons and sprouting fiber bundles.

Ventral lateral nucleus (thalamus, embryonic) – Situated caudal to the *ventral anterior nucleus, this component of the thalamic *ventral complex is the target of input from the *superior cerebellar peduncle and the *red nucleus. Its somatotopically organized fibers terminate in the motor cortex and adjacent areas. The ventral lateral nucleus may be the principal relay from the *cerebellum to the *neocortex. The nucleus is recognizable during the late first trimester as a developing structure with putative migrating and sojourning neurons and sprouting fiber bundles.

Ventral posterolateral nucleus (thalamus, embryonic) – Situated caudal to the *ventral lateral nucleus, this region of the thalamic *ventral complex is the target of fibers of the *medial lemniscus that originate in the *cuneate nucleus and the *gracile nucleus and convey somatosensory information from the trunk and the extremities. Its efferents form the somesthetic radiation that terminates in a precise topographic order in the medial part of the *postcentral gyrus. The ventral posterolateral nucleus is the principal thalamic relay of somesthetic input from the trunk and limbs to the *neocortex. The nucleus is recognizable during the late first trimester as a developing structure with putative migrating and sojourning neurons and sprouting fiber bundles.

Ventral posteromedial nucleus (thalamus, embryonic) – Situated between the *ventral posterolateral nucleus and the *centromedian nucleus, this nucleus receives afferents from the *trigeminal sensory nuclei and the *parabrachial nucleus that convey sensory information from the face, the tongue, the oral cavity, and the neck. The efferents of this nucleus terminate in a precise topographic order in the lateral part of the *postcentral gyrus. The ventral posteromedial nucleus is the principal thalamic relay of somatosensory and gustatory input from the neck, head, and mouth to the *neocortex. The nucleus is recognizable during the late first trimester as a developing structure with putative migrating and sojourning neurons and sprouting fiber bundles.

Ventral rhombic lip – *see* **Precerebellar NEP**

Ventral tegmental area (embryonic) – Medial area flanking the *substantia nigra and containing a high concentration of dopaminergic neurons, much like the

substantia nigra, pars compacta. It is present as a compact cell mass in the mesencephalon of late first trimester fetuses.

Ventricles – *See* **Supraventricles**

Ventromedial nucleus (hypothalamus) – Large spherical nucleus that flanks the third ventricle and is surrounded by a fibrous shell. It has reciprocal connections with the *amygdala, the *bed nucleus of the stria terminalis, the *septum, and the *subiculum. It has been implicated in motivational functions related to feeding and sexual behavior.

Vermis – *See* **Cerebellum (vermis)**

Vestibular nuclear complex – A large area in the dorsal medulla, composed of the *medial, the *lateral, the *superior, and the *inferior vestibular nuclei. These nuclei get primary sensory input from the vestibular ganglion; their efferents join the *medial longitudinal fasciculus and form the vestibulospinal tract.

Visual radiation (embryonic) – Thalamocortical fibers that originate in the *lateral geniculate body and terminate in the striate cortex of the *occipital lobe. The identification of *Meyer's loop at GW11 suggests that this tract may reach the occipital lobe by the end of the first trimester.

W

White matter – General term for extensive regions in the brain and spinal cord composed of myelinated fiber tracts but few or no neuronal cell bodies. In histological preparations with myelin stains, the white matter appears black. In laminated brain regions, as in the *cerebral cortex, the white matter is called the medullary layer.

Z

Zygomatic bone – A facial bone in the cheek bone.

Zona incerta (embryonic) – Region in the *subthalamus with uncertain boundaries with *Forel's fields. It is a prominent area in the late first trimester *diencephalon.

An asterisk in front of a term indicates that it is a separate entry in the Glossary with additional information. Terms referring to transient developmental structures are underlined.

