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Hypothalamus

The **hypothalamus** is a portion of the brain that contains a number of small nuclei with a variety of functions. One of the most important functions of the hypothalamus is to link the nervous system to the endocrine system via the pituitary gland. The hypothalamus is located below the thalamus and is part of the limbic system.^[1] In the terminology of neuroanatomy, it forms the ventral part of the diencephalon. All vertebrate brains contain a hypothalamus. In humans, it is the size of an almond. The hypothalamus is responsible for the regulation of certain metabolic processes and other activities of the autonomic nervous system. It synthesizes and secretes certain neurohormones, called releasing hormones or hypothalamic hormones, and these in turn stimulate or inhibit the secretion of hormones from the pituitary gland. The hypothalamus controls body temperature, hunger, important aspects of parenting and attachment behaviours, thirst,^[2] fatigue, sleep, and circadian rhythms. The hypothalamus derives its name from Greek ὑπὀ, under and θάλαμος, chamber.

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Structure

The hypothalamus is a brain structure made up of distinct nuclei as well as less anatomically distinct areas. It is found in all vertebrate nervous systems. In mammals, <u>magnocellular neurosecretory cells</u> in the <u>paraventricular nucleus</u> and the <u>supraoptic nucleus</u> of the hypothalamus produce <u>neurohypophysial hormones</u>, <u>oxytocin</u> and <u>vasopressin</u>. These hormones are released into the blood in the <u>posterior pituitary</u>.^[3] Much smaller <u>parvocellular neurosecretory cells</u>, neurons of the paraventricular nucleus, release <u>corticotropin-releasing hormone</u> and other hormones into the <u>hypophyseal portal system</u>, where these hormones diffuse to the anterior pituitary.

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Anatomical terms of neuroanatomy

Nuclei

The hypothalamic nuclei include the following:^{[4][5][6]}



Human hypothalamus (shown in red)

Region	Area	Nucleus	Function ^[7]
Anterior	Preoptic	Preoptic nucleus	Thermoregulation
	Medial	Medial preoptic nucleus	 Regulates the release of gonadotropic hormones from the adenohypophysis Contains the <u>sexually dimorphic nucleus</u>, which releases GnRH, differential development between sexes is based upon in utero testosterone levels Thermoregulation^[8]
		Supraoptic nucleus	 <u>Vasopressin</u> release <u>Oxytocin</u> release
		Paraventricular nucleus	 thyrotropin-releasing hormone release corticotropin-releasing hormone release oxytocin release vasopressin release somatostatin release
		Anterior hypothalamic nucleus	 thermoregulation panting sweating thyrotropin inhibition
		Suprachiasmatic nucleus	 Circadian rhythms
	Lateral	Lateral nucleus	See Lateral hypothalamus § Function – primary source of orexin neurons that project throughout the brain and spinal cord
Tuberal	Medial	Dorsomedial hypothalamic nucleus	 blood pressure heart rate GI stimulation
		Ventromedial nucleus	 satiety neuroendocrine control
		Arcuate nucleus	 Growth hormone-releasing hormone (GHRH) feeding Dopamine-mediated prolactin inhibition
	Lateral	Lateral nucleus	See Lateral hypothalamus § Function – primary source of orexin neurons that project throughout the brain and spinal cord
		Lateral tuberal nuclei	
Posterior	Medial	Mammillary nuclei (part of mammillary bodies)	■ memory

List of nuclei, their functions, and the neurotransmitters, neuropeptides, or hormones that they utilize

	Posterior nucleus	 Increase <u>blood pressure</u> <u>pupillary</u> dilation <u>shivering</u> <u>vasopressin</u> release
	Lateral nucleus	See Lateral hypothalamus <u>§</u> Function – primary source of orexin neurons that project throughout the brain and spinal cord
Lateral	Tuberomammillary nucleus ^[9]	 arousal (wakefulness and attention) feeding and <u>energy balance</u> learning memory sleep

See also: ventrolateral preoptic nucleus, periventricular nucleus.





A cross section of the Hypothalamic monkey hypothalamus Symbols: AC displays 2 of the major commissure hypothalamic nuclei on preoptic nuclei either side of the fluid-suprachiasma filled 3rd ventricle. OC: optic ch

Symbols: AC: anterior commissure PO: SC: preoptic nucleus suprachiasmatic nucleus OC: optic chiasma TC: tuber cinereum AP: pituitary IN: anterior infundibulum PP: posterior pituitary ME: median eminence AH: anterior hypothalamic nucleus SO: supraoptic nucleus TH: thalamus PV: paraventricular nucleus (not to be confused with periventricular nucleus, which is not shown) DM: dorsomedial nucleus VM: ventromedial nucleus AR: arcuate nucleus (associated with periventricular nucleus, which is not shown) LT: PN: lateral nucleus posterior nucleus MB: mamillary body

nuclei Hypothalamic nuclei on nterior one side of the PO: hypothalamus, shown in SC: a 3-D computer ucleus reconstruction^[10]

Connections

The hypothalamus is highly interconnected with other parts of the <u>central nervous system</u>, in particular the brainstem and its <u>reticular formation</u>. As part of the <u>limbic system</u>, it has connections to other limbic structures including the <u>amygdala</u> and <u>septum</u>, and is also connected with areas of the <u>autonomous nervous system</u>.

The hypothalamus receives many inputs from the brainstem, the most notable from the nucleus of the solitary tract, the

locus coeruleus, and the ventrolateral medulla.

Most nerve fibres within the hypothalamus run in two ways (bidirectional).

- Projections to areas <u>caudal</u> to the hypothalamus go through the <u>medial forebrain bundle</u>, the <u>mammillotegmental</u> tract and the <u>dorsal longitudinal fasciculus</u>.
- Projections to areas rostral to the hypothalamus are carried by the <u>mammillothalamic tract</u>, the <u>fornix</u> and <u>terminal</u> <u>stria</u>.
- Projections to areas of the sympathetic motor system (lateral horn spinal segments T1-L2/L3) are carried by the hypothalamospinal tract and they activate the sympathetic motor pathway.

Sexual dimorphism

Several hypothalamic nuclei are <u>sexually dimorphic</u>; i.e., there are clear differences in both structure and function between males and females.^[11] Some differences are apparent even in gross neuroanatomy: most notable is the <u>sexually</u> <u>dimorphic nucleus</u> within the <u>preoptic area</u>,^[11] in which the differences are subtle changes in the connectivity and chemical sensitivity of particular sets of neurons. The importance of these changes can be recognized by functional differences between males and females. For instance, males of most species prefer the odor and appearance of females over males, which is instrumental in stimulating male sexual behavior. If the sexually dimorphic nucleus is lesioned, this preference for females by males diminishes. Also, the pattern of secretion of <u>growth hormone</u> is sexually dimorphic;^[12] this is why in many species, adult males are visibly distinguishable from females.

Responsiveness to ovarian steroids

Other striking functional dimorphisms are in the behavioral responses to <u>ovarian steroids</u> of the adult. Males and females respond to ovarian steroids in different ways, partly because the expression of estrogen-sensitive neurons in the hypothalamus is sexually dimorphic; i.e., estrogen receptors are expressed in different sets of neurons.

Estrogen and progesterone can influence gene expression in particular neurons or induce changes in <u>cell membrane</u> potential and <u>kinase</u> activation, leading to diverse non-genomic cellular functions. Estrogen and progesterone bind to their cognate <u>nuclear hormone receptors</u>, which translocate to the cell nucleus and interact with regions of DNA known as <u>hormone response elements</u> (HREs) or get tethered to another <u>transcription factor</u>'s binding site. <u>Estrogen receptor</u> (ER) has been shown to transactivate other transcription factors in this manner, despite the absence of an <u>estrogen response element</u> (ERE) in the proximal promoter region of the gene. In general, ERs and <u>progesterone receptors</u> (PRs) are gene activators, with increased mRNA and subsequent protein synthesis following hormone exposure.

Male and female brains differ in the distribution of estrogen receptors, and this difference is an irreversible consequence of neonatal steroid exposure. Estrogen receptors (and progesterone receptors) are found mainly in neurons in the anterior and mediobasal hypothalamus, notably:

- the preoptic area (where LHRH neurons are located, regulating dopamine responses and maternal behavior;^[13]
- the periventricular nucleus where somatostatin neurons are located, regulating stress levels;[14]
- the ventromedial hypothalamus which regulates hunger and sexual arousal.

Development

In neonatal life, gonadal steroids influence the development of the neuroendocrine hypothalamus. For instance, they determine the ability of females to exhibit a normal reproductive cycle, and of males and females to display appropriate reproductive behaviors in adult life.