

Dorsal raphe nucleus

The dorsal raphe nucleus (DRN) is a heterogeneous [brainstem nucleus](#) located in the [midbrain](#) and [pons](#). Via widespread projections, which target a multitude of brain areas, its neurons utilize many transmitters to control various physiological functions, including learning, memory and affect.

Previous studies have demonstrated that [Serotonin](#) type 3 (5-HT₃) receptors are expressed in either or both the [substantia nigra](#) (SN) and the dorsal raphe nucleus (DRN) in humans, marmosets, rats and Syrian hamsters.

Belliveau et al. quantified the distribution of 5-HT₃ receptors across these regions in the adult rat. Fluorescent [immunohistochemistry](#) was performed on sections of rat brain covering the entire rostro-caudal extent of the SN and DRN with antibodies specific to the 5-HT_{3A} receptor subunit, as well as others targeting the monoaminergic markers tyrosine hydroxylase (TH) and the 5-HT transporter (SERT). The number of 5-HT_{3A} receptor-positive, TH-positive ($n = 28,428 \pm 888$, Gundersen's $m = 1$ coefficient of error [CE] = 0.05) and SERT-positive ($n = 12,852 \pm 462$, CE = 0.06) cells were estimated in both the SN and the DRN using stereology. We found that 5-HT_{3A} receptor-positive cells are present in the SNr ($n = 1,250 \pm 64$, CE = 0.24), but they did not co-localise with TH-positive cells, nor were they present in the SNc. In contrast, no 5-HT_{3A} receptor-positive cells were found in the DRN. These results support the presence of 5-HT₃ receptors in the SN, but not in the DRN, and do not support their expression on monoaminergic cells within these two brain areas ¹⁾.

The other main brain entry of the information related to fluid and cardiovascular balance are the [lamina terminalis](#) (LT) and one of the sensory circumventricular organs (CVOs), the [area postrema](#) (AP). The LT, consisting of the [median preoptic nucleus](#) (MnPO) and the other two sensory CVOs—i.e., [subfornical organ](#) (SFO) and [organum vasculosum](#) of the lamina terminalis (OVLT)—is recognized as a site in the brain that is crucial for the physiological regulation of hydroelectrolyte balance. The SFO and OVLT lack a blood-brain barrier and contain cells that are sensitive to humoral signals, such as changes in plasma and cerebrospinal fluid sodium concentration (Vivas et al. 1990), [osmolality](#) (Sladek and Johnson 1983), and [angiotensin II](#) (ANG II) levels (Ferguson and Bains 1997; Simpson et al. 1978). Such unique features make the SFO and OVLT key brain regions for sensing the status of the body fluids and electrolytes. Humoral and neural signals that arrive to the two main brain entries—that is, the CVOs of the LT and within the [hindbrain](#) the AP-NTS—activate a central circuit that includes integrative areas such as the MnPO, the paraventricular (PVN), the supraoptic (SON), [lateral parabrachial nucleus](#) (LPBN), [dorsal raphe nucleus](#) (DRN), and neurochemical systems such as the angiotensinergic, vasopressinergic, oxytocinergic (OT), and serotonergic (5-HT) systems.

Once these signals act on the above-mentioned neurochemical networks, they trigger appropriate sympathetic, endocrine, and behavioral responses. Therefore, after a body fluid deficit, water and [sodium](#) intake and excretion need to be controlled to minimize disturbances of hydromineral homeostasis. In this context, [hypovolemia](#) and [hyponatremia](#) induced by body fluid depletion stimulate central and peripheral osmo-sodium receptors, taste receptors, volume and arterial/cardiopulmonary baroreceptors, and the renin-angiotensin system (RAS). This latter system, for example, acts mainly through the sensory CVOs and/or the AP to activate brain neural pathways that elevate BP, release vasopressin and aldosterone (ALDO), increase renal sympathetic nerve activity, and increase the ingestion of water and sodium ²⁾

1)

Belliveau S, Kang W, Bovaird S, Hamadjida A, Bédard D, Dancause N, Stroh T, Huot P. Stereological investigation of 5-HT₃ receptors in the substantia nigra and dorsal raphe nucleus in the rat. *J Chem Neuroanat*. 2020 Nov 4:101881. doi: 10.1016/j.jchemneu.2020.101881. Epub ahead of print. PMID: 33160048.

²⁾

Vivas L, Godino A, Dalmaso C, Caeiro XE, Macchione AF, Cambiasso MJ. Neurochemical Circuits Subserving Fluid Balance and Baroreflex: A Role for Serotonin, Oxytocin, and Gonadal Steroids. In: De Luca LA Jr, Menani JV, Johnson AK, editors. *Neurobiology of Body Fluid Homeostasis: Transduction and Integration*. Boca Raton (FL): CRC Press/Taylor & Francis; 2014. Chapter 9. PubMed PMID: 24829993.

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