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Cerebral vasodilator

mechanisms in intracranial

hypertension. Cerebral

vasodilators that are active in

the central nervous system include the vasoconstrictors adenosine, vasopressin, substance P, calcitonin gene-related peptide, and the nitric oxide (NO) donor nitroglycerin, and the NO synthase (NOS) inhibitors NG-monomethyl L-arginine (L-NMMA) and NG-nitro-L-arginine (L-NNA).

Cerebrospinal fluid (CSF) levels of these compounds are elevated in conditions of increased intracranial pressure.

Because they are in direct competition with other endogenous NO donors for

binding to the same receptor, the study of the effects of these compounds in the CSF, and the interpretation of their effects in the central nervous system, depends on knowledge of the regulation of the production and release of these compounds. L-NMMA is a potent inhibitor of NOS I and II, but it has no direct effect on L-arginine transport in brain microvascular endothelial cells in vitro. L-NMMA and L-NNA are competitive inhibitors of NOS III, with inhibitory constants that vary from 1 to

100 microM depending on the assay system. NG-methyl-L-arginine inhibits L-arginine uptake by cultured rat cortical astrocytes in a dose-dependent manner. However, astrocytes do not produce NG

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