



## Drugs as Tools in Neurotransmitter Research (Neuromethods)

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Serendipitous discovery has played an important role in the identification of physiological mechanisms in central and peripheral neurotransmission. In many cases, the subsequent follow-up of these discoveries has been substantially aided by previously identified natural substances that interfered with these mechanisms. The site of neuromuscular blockade produced by curare extracts, for example, was shown in the 1850s by Claude Bernard, and Sir Henry H. Dale in 1906 discovered the reversal of the pressor effects of adrenal extracts by ergot alkaloids. These findings were the first clues that led eventually to the definition of nicotinic cholinergic receptors and the present classification of adrenergic receptors. In the 1950s many discoveries, including the identification of the catecholamines and 5-hydroxytryptamine in the central nervous system, followed by the development of specific sensitive biochemical and histochemical methods and the introduction of modern psychotropic drugs led to our increased understanding of chemical transmission and the formulation of theories that attempt to explain the causes of some mental disorders. The finding that hydroxylated derivatives of noradrenaline and 5-hydroxytryptamine are readily taken up by catecholaminergic or serotonergic neurons, and that this is followed by poisoning and death of the cell, encouraged the use of these compounds for the mapping of central and peripheral catecholamine or 5-hydroxytryptamine systems, and the investigation of the relevance of these pathways to receptor regulation or behavioral responses.

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