

Poster presentation

Cognitive disorders in 6-hydroxydopamine-induced rat model of Parkinson's disease

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Background

Parkinson's disease is a human neurodegenerative disorder which is mainly characterized by a massive and progressive degeneration of the dopaminergic neurons in the substantia nigra (SN). The most widely used animal models of Parkinson's disease involve intracranial infusion of the neurotoxin 6-hydroxydopamine (6-OHDA) directly into the ascending dopaminergic forebrain bundle, thereby, inducing severe dopaminergic neuronal degeneration associated with profound deficits in feeding, drinking, sensorimotor and learning functions [1-4]. The aim of the present work was to study the effects of right-unilateral 6-OHDA lesions of the ventral tegmental area (VTA) or substantia nigra pars reticulata (SNr) on learning and memory processes evidenced by means of Y-maze task and shuttle-box task, respectively. We also examined the effect of nicotine treatment on the 6-OHDA lesioned rats. Our data suggest that a correlation exist between VTA, SNr and nAChRS and expression of cognitive capacities.

Materials and methods

Male Wistar rats were subjected to right-unilateral 6-hydroxydopamine (6-OHDA) (8µg/4µl) lesions of the ventral tegmental area (VTA) or substantia nigra pars reticulata (SNr), or were sham lesioned, and nicotine treatment and their ability to acquire the operant task was studied by means of Y-maze task and shuttle-box task, respectively. For the lesioning of the SN the following coordinates were used: 5.5 mm.

Posterior to bregma; 2.0 mm lateral to the midline; 7.4 mm ventral to the surface of the cortex. For lesioning the VTA, the following coordinates were used: 5.6 mm.

Posterior to bregma; 0.5 mm lateral to the midline; 7.6 mm ventral to the surface of the cortex. The sham-operated rats were injected with saline. Learning and memory tests were began 2 weeks after the operation.

Results

Lesions of both VTA and SNr resulted in an impairment of both conditioned avoidance response and crossing latency tested by means of shuttle-box task, suggesting significant effects of long-term memory. 6-OHDA significantly decreased spontaneous alternation in Y-maze task, suggesting effects on spatial memory, especially on short-term memory. A low dose of nicotine (0.3 mg/kg b.w., i.p.), a specific nicotinic acetylcholine receptors (nAChRS), administrated 4 consecutively days attenuated the impairment of learning and memory processes in 6-OHDA lesioned rats.

Conclusions

These data suggest that VTA, SN and nAChRS have a facilitator role in learning and memory processes. Therefore, the integrity of these nervous areas may be necessary for processing and storage of information.

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