

Original Paper

Effect of donepezil hydrochloride on reference and working memory impairment after bilateral electrical lesion of nucleus basalis magnocellularis in rats model of Alzheimer disease

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Abstract

Background and Objective: Dysfunction and loss of basal forebrain cholinergic neurons and their cortical projections are the earliest pathological events in the pathogenesis of Alzheimer disease (AD). This study was done to evaluate the effect of donepezil hydrochloride on reference and working memory caused by mutual electrical lesion of the nucleus basalis magnocellularis (NBM) in animal model of AD.

Methods: In this experimental study, 56 adult male Wistar rats were allocated into 8 groups (n=7) including: control (intact), NBM lesion group, which received electrically-induced lesion (0.5 mA, 3s) in NBM, Sham group (the electrode was impaled into the NBM with no lesion), donepezil groups (lesion + 0.1, 1, 5, 10 mg/kg/bw of donepezil hydrochloride) and vehicle group (NBM lesion + saline). Acquisition and retention testing were done by using an eight-radial arm maze, in which, the patterns of arm entries in each group were recorded for calculating correct choice, working memory errors, reference memory error and latency.

Results: The spatial learning of animals in the lesion of NBM group significantly reduced compared to controls (P<0.05). Moreover, no effect on spatial learning was seen in the sham group compared with the lesion group. The post-lesion treatment with donepezil hydrochloride in a dose-dependent manner improved the parameters of spatial memory errors in the acquisition and retention tasks in comparison with the lesion group (P<0.05).

Conclusion: Treatment with donepezil hydrochloride, dose-dependently improves cognitive impairment induced by the destruction of the nucleus basalis magnocellularis.

Keywords: Alzheimer disease, Nucleus Basalis Magnocellularis, Memory, Donepezil hydrochloride, Rat

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