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ACCEPTED MANUSCRIPT

Toxic Influence of Paraquat on Hippocampal

Neurogenesis in Adult Mice

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Abstract: Paraquat, a fast-acting non-selective contact herbicide, is considered an etiological

factor related to Parkinson's disease. This study investigated its effects on hippocampal

neurogenesis and cognition in adult mice as well as possible mechanisms for the effects. We

administered paraquat (1.25 mg/kg, intraperitoneal injection, i.p.) and an equal volume of normal

saline for 3 weeks to adult male C57BL/6J mice. The results showed that hippocampus-dependent

spatial learning and memory was significantly impaired in paraquat-treated mice. Moreover,

paraquat administration inhibited the proliferation of neural progenitor cells, and impaired the

survival and altered the fate decision of newly generated cells in the hippocampus. The expression

levels of caspase-3 and glial fibrillary acidic protein were significantly higher in paraquat-treated

mice than in control mice. Interestingly, paraquat reduced the phosphorylation of Akt, but did not

affect the total amount of Akt. In conclusion, our findings suggest that paraquat negatively

affected adult hippocampal neurogenesis and cognition function.

Key Words: paraquat; learning; memory; neurogenesis; phosphorylation

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