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## The neuroprotective effect of Tocotrienol in chronic cerebral hypoperfusion-induced neurodegeneration in rats

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**Introduction:** Reduced cerebral blood flow (CBF) is associated with aging and neurodegenerative disorders. CBF-induced neurodegeneration is related with the formation of reactive oxygen species (ROS), which is fatal to neurons at high concentration. To study the neuropathological consequences of a reduced CBF, a similar condition has been created in rats by common carotid artery occlusion (2 vessel occlusion, 2VO). Since vitamin E is known to be a potent antioxidant, the present study, therefore, was designed to assess the effects of vitamin E as an antioxidant and neuroprotective agent in 2VO rat model.

**Materials & Methods:** After acclimatization, twenty four Sprague Dawley rats weighing 200-250 g were equally divided into three groups. Group A: Sham control, Group B: 2VO and Group C: 2VO+E (treated daily with Vit E, 100 mg/kg, orally following 2VO). On the 8th week, all the rats were euthanized and the hippocampi were isolated. Viable neuronal cell count in the hippocampal CA-1 region was estimated. The Isoprostane F2 (Iso-F2) levels were also measured in the brain homogenates to quantify the oxidative stress levels.

**Results:** There was significant difference in neuronal cell death in 2VO group as compared to sham group. In 2VO+E rats, the viable neuronal cell count of the hippocampal CA-1 region was significantly higher ( $p < 0.05$ ) as compared to the 2VO group. Moreover, Iso-F2 levels in 2VO group was significantly higher ( $p < 0.05$ ) as compared to 2VO+E group, implying high oxidative stress in 2VO group and reduction of oxidative stress levels in 2VO+E group.

**Conclusion:** This study clearly demonstrates the effectiveness of Vit E as a neuroprotective and antioxidant agent in chronic cerebral hypoperfusion induced-neurodegeneration in rats.

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## Managing a global regulatory affairs strategy

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Having a solid, global regulatory affairs strategy is essential to efficient and effective approval and commercialization of pharmaceutical products. This presentation will review similar and divergent regulatory strategies in key areas around the world and provide practical advice on managing a timely and high quality program to achieve success.

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