学位論文要約 Extended Summary in Lieu of the Full Text of a Doctoral Thesis

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学位論文題目:

Endothelium-dependent vasodilation in the cerebral arterioles of rats

deteriorates during acute hyperglycemia and then is restored by reducing

Thesis Title the glucose level.

学位論文要約:

Summary of Thesis

Purpose: Acute hyperglycemia in patients with traumatic brain injury correlates with a poor neurolo gical outcome. We investigated the endothelium function of rat cerebral arterioles during acute hyperglycemia and after reducing the glucose levels using insulin. We also examined whether or not o xidative stress was involved in the cerebral arteriole response to acute hyperglycemia.

Methods: In isoflurane-anesthetized, mechanically ventilated rats, we used closed cranial window pr eparation to measure the changes in the pial arteriolar diameter following the topical application of acetylcholine (ACh) or adenosine. We examined the pial arteriolar vasodilator response before hyp erglycemia, during hyperglycemia, and after reducing the glucose level using insulin. After intraven ous pretreatment with an NADPH oxidase inhibitor (apocynin or diphenylene iodonium), we reexa mined the pial arteriolar vasodilator response following the topical application of ACh.

Results: Under control conditions, the topical application of ACh dose-dependently dilated the cereb ral arterioles. The vasodilatory responses to topical ACh were impaired during hyperglycemia and i mproved after the administration of insulin. The vasodilatory responses to topical adenosine were n ot affected by the glucose levels. In the apocynin or diphenylene iodonium pretreatment group, the topical application of ACh dilated the cerebral arterioles during hyperglycemia.

Conclusion: Acute hyperglycemia induces a dysfunction of the endothelium-dependent vasodilation of rat cerebral arterioles. The dysfunction can be reversed by improving the acute hyperglycemia and it can be prevented entirely by the administration of NADPH oxidase inhibitors. These results could suggest that controlling the glucose levels work protectivity to endothelium function of cerebral arterioles.