

## P23 Neuroprotective Effects of KC0244, a Glycine Site Antagonist, in a Rat Model of Transient Focal Ischemia

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Antagonists acting at the glycine site of the NMDA receptor have been gaining safer alternatives for stroke therapy because they have few adverse effects compared with competitive and noncompetitive NMDA antagonists. Therefore, the neuroprotective novel glycine site antagonist KC0244 were evaluated in a rat model of transient focal ischemia in comparison with GV150526A in a developmental phase. Middle cerebral artery occlusion was produced by insertion of a silicone-coated 4-0 nylon monofilament to the ophthalmic artery in male Sprague-Dawley rats under isoflurane anesthesia. After 90 or 120 min of occlusion, the filament was retracted and the ischemic tissue reperused. In 90-min MCAO model, GV150526A was administered 30 min before MCAO or immediately after MCAO. In 120-min MCAO model, KC0244 or GV150526A (10 mg/kg, i.p.) was administered 1 hr before MCAO or immediately after MCAO. Infarct volume was measured 24 hr after MCAO using the 2,3,5-triphenylthiazine chloride staining method. In 90-min MCAO model, treatments with GV150526A significantly reduce infarct volume although they tended to slightly reduce cortical infarct volume approximately 19% compared with the nontreated group. In 120-min MCAO model, treatments with GV150526A did not either significantly reduce infarct volume although they tended to reduce total infarct volume by approximately 16% compared with the vehicle-treated group. However, 1-hr preischemic and immediate treatments with KC0244 reduced total infarct volume by 39 and 30% (corrected total infarct volume by 44 and 32%), respectively, compared with the vehicle-treated control group. The results suggest that KC0244 can provide neuroprotection against transient focal ischemic damage with greater *in vivo* potency than GV150526A.