

In the early stages of the war, when Grafton's regiment was ordered into western Virginia, where the fighting was heaviest, Grant personally went to Boston to pick up this detachment at its starting point in New York city. 8, 1818, gpspr

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you do it?.A new family

of quinolinic acid

inhibitors. Synthesis and

biological activity. A new

class of glutamate

antagonists has been

developed based on our

previous work demonstrating that quinolinic acid (QUIN) is neurotoxic and that 2-methyl-QUIN (2-MeQUIN) is not. In the present study, we have synthesized a series of 2,4-disubstituted quinolinic acids and attempted to elucidate the structure-activity relationship (SAR) in

order to determine the structural features required for neurotoxicity and to determine which of the amino acid conformation available to the drug is critical in determining activity. New derivatives were evaluated in vitro for neurotoxicity, platelet activation, and excitotoxicity. Four

compounds were found to be neurotoxic, as evidenced by their ability to reduce the mitochondrial membrane potential (MMP) in cultured hippocampal neurons. The most potent of these compounds was 2,4-dihydroxy-3-methoxy-quinolinic acid (2,4-DEO-3-MeQUIN).

Among the nonneurotoxic compounds, the one with the greatest inhibitory effect on platelet aggregation in vitro was also the most active in reducing quinolinic acid-induced excitotoxicity. Although the position of the amino acid side chain on the quinolinic acid molecule is not

critical for neurotoxicity,
replacement of the ethyl
side chain in 2-MeQUIN
with a methyl group
increased the inhibition
of platelet aggregation
and excitotoxicity. This
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