

论文

疟疾防治药物的研究——X V. 双-(2,4-二氨基喹唑啉-6-取代氨基甲基)芳香类化合物的合成及其抗疟作用

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摘要:

关键词: 抗疟作用 二氢叶酸还原酶抑制剂 双-(2,4-二氨基喹唑啉-6-取代氨基甲基)芳香类化合物

STUDIES ON ANTIMALARIALS——X V. SYNTHESIS AND ANTIMALARIAL ACTIVITIES OF SOME BIS(2, 4-DIAMINOQUINAZOL-6-YL-SUBSTITUTED AMINOMETHYL) AROMATIC DERIVATIVES

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Abstract:

A series of bis(2,4-diamino-quinazol-6-yl-substituted aminomethyl) aromatic derivatives were synthesized. Compounds II_{1~5} were synthesized by two ways. In the first, 2,4,6-triaminoquinazoline was condensed with the corresponding aryl dialdehydes to form Schiff bases which were reduced with sodium borohydride; the second used a new method in which 2,4,6-triaminoquinazoline was directly condensed with the corresponding aralkyl dihalides to afford the same products. The derivatives II_{6~16} were prepared by formylation, nitrosation or methylation respectively. The inhibition activities of the title compounds on dihydrofolate reductase in rat liver were assayed. The activities of formylated or nitrosated products were 6-9 times, and of methylated products were twofold as active as that of the parent compounds. II₆, II₉, II₁₀ and II₁₄ were nearly as potent as pyrimethamine. Primary antimalarial screening in mice showed that no one possessed significant activity.

Keywords: Inhibitor for dihydrofolate reductase Bis (2, 4-diaminoquinazol-6-yl-suhstituted aminomethyl)-aromatic derivatives. Antimalarial action

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