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论文

S-9-(2,3-二羟丙基)腺嘌呤类似物的合成及其对S-腺苷-L-高半胱氨酸水解酶的抑制活性

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摘要:

为了寻找高效、低毒的抗病毒剂,本文通过用腺嘌呤及嘧啶碱基与3-氯-2-甲基丙烯缩合合成6个DHPA类似物的中间 体5a,5b,6~8a和8b。应用OsO,催化,在N-甲基吗琳-N-氧化物氧化下对烯键进行邻位双羟化,合成了5个DHPA类似 物1~4a,b。对化合物5a,5b,6~8a,8b的 1 HNMR数据进行了初步总结。对4个DHPA类似物测定了它们对S-腺苷-L-高半胱氨酸水解酶(SAH)的抑制活性,其中化合物1的 IC_{50} 为1.1 $mmol\cdot L^{-1}$,其余化合物均无抑制活性。

关键词: 开链核苷 DHPA类似物 SAH水解酶 抗病毒剂

SYNTHESIS OF S-9-(2,3-DIHYDROXYPROPYL)ADENINE (DHPA) ANALOGS AND THEIR ▶ 开链核苷 INHIBITION OF S-ADENOSYL-L-HOMOCYSTEINE (SAH) HYDROLASE

GX Wang; L Wang; ZZ Zhao; T Zhang * and PZ Tao

Abstract:

In order to search for new antiviral agents with high potency and low toxicity, eleven new acyclonucleosides were synthesized. Nucleic acid bases were condensed with 3-chloro-2-methylpropene to give $5\sim 8a.b$, which were oxidized by N-methylmorpholine N-oxide in the presence of OsO₄ to give vicinal dihydroxy acyclonucleosides 1~4a.b. Four DHPA analogs have been tested for the inhibitory activities on s adenosyl L homocysteine hydrolase (SAH). Only compound 1 showed some enzyme inhibitory effect with IC 50 of 1.1 mmol·L⁻¹

Keywords: Acyclonucleosides Antiviral agents S-adenosyl-L-homocysteine hydrolase

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