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论文

3-苯基-4(1H)喹啉酮羟基衍生物的合成及其抗骨质疏松活性

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摘要:

目的设计并合成7-羟基-3-(取代)苯基-4(1H)喹啉酮化合物及其相应的5-羟基-3-(取代)苯基-4(1H)喹啉酮化合物并考察其抗骨质疏松活性。方法以间氨基酚为原料,选择改良的Gould-Jacobs反应路线同时合成7-羟基-3-(取代)苯基-4(1H)喹啉酮4个($A_{1\sim4}$)及其相应的5-羟基-3-(取代)苯基-4(1H)喹啉酮4个($B_{1\sim4}$),通过骨细胞筛选实验以及羟磷灰石吸附实验分别考察其促骨形成作用和趋骨性。结果共合成了新化合物8个($A_{1\sim4}$, $B_{1\sim4}$),其结构经IR,MS,1HNMR和元素分析确证。骨细胞筛选实验结果表明,B₃有促骨形成作用,但作用弱于芒柄花黄素;羟磷灰石吸附实验结果表明,羟磷灰石对B₁,B₂和B₄有一定的吸附,其中对B₁和B₂的吸附作用强于四环素。结论 5-羟基-3-(取代)苯基-4(1H)喹啉酮化合物有促骨形成作用并具有一定的趋骨活性。

关键词: 3-苯基-4(1H)喹啉酮衍生物 抗骨质疏松 促骨形成 趋骨性 羟磷灰石

SYNTHESIS AND ANTIOSTEOPOROSIS ACTIVITY OF 3-PHENYL-4(1H) QUINOLINONE DERIVATIVES

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Abstract:

AIMTo design and synthesize 7-hydroxy-3-(substituted) phenyl-4(1H) quinolinone and 5-hydroxy-3-(substituted) phenyl-4(1H) quinolinone and to study the antiosteoporosis activity of these compounds. METHODSThe compounds of 7-hydroxy and 5-hydroxy were prepared simultaneously by improved Gould-Jacobs reaction and resorcin was used as starting material. Eight compounds ($A_{1\sim 4}$, $B_{1\sim 4}$) were synthesized and their chemical structures were determined by IR, MS, ¹HNMR and elemental analysis. Their osteoplastic activity was evaluated by the absorptive test to Ca^{2+} of hydroxyapatite crystal. RESULTSCompounds ($A_{1\sim 4}$, $B_{1\sim 4}$) are new compounds. Compounds B_1 and B_2 showed antiosteoporosis activities and was stronger than the control drug of tetracycline. CONCLUSIONCompounds B_1 and B_2 are worthy to be intensively studied.

Keywords: antiosteoporosis osteoplastic activity osteocyte chemotaxis hydroxyapatite 3-phenyl-4 (1H) quinolinone derivatives

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