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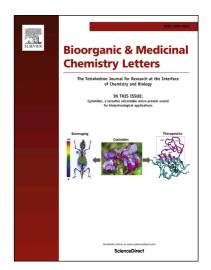
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ACCEPTED MANUSCRIPT

Isosteric ribavirin analogues: synthesis and antiviral activities.

Nikolay I. Zhurilo^a, Mikhail V. Chudinov^{a†}, Andrey V. Matveev^a, Olga S. Smirnova^b, Irina D. Konstantinova^b, Anatoly I. Miroshnikov^b, Alexander N. Prutkov^c, Lyubov E. Grebenkina^c, Natalya V. Pulkova^c, Vitaly I. Shvets^a

Abstract

The novel isosteric ribavirin analogues were synthesized by two different ways. Some of them showed significant antiviral action against hepatitis C virus (HCV), herpes simplex (HCV-1) and influenza A virus comparable to that of ribavirin itself. The data obtained confirm the proposed theory of the ribavirin possible antiviral activity mechanism related with bioisosterism.

Ribavirin (Virazole, 1- β -D-ribofuranosyl-1,2,4-triazole-3-carboxamide) is a nucleoside analogue with a broad antiviral activity spectrum. It is active against influenzia virus, HCV, RSV, HSV etc. $^{1-6}$.

Ribavirin action mechanism is still not clarified, however, an existing hypothesis assume the heterocyclic base of ribavirin, 1,2,4-triazole-3-carboxamide (TKA), as a mimic of guanosine purine cycle. This steric similarity allows to ribavirin molecule to incorporate into the viral genome and cause irreversible mutations⁷.

^a Lomonosov Institute of Fine Chemical Tehnologies, Moscow Technological University, Vernadskogo Pr. 78, 119454, Moscow, Russia.

^b Shemyakin and Ovchinnikov Institute of Bioorganic Chemistry, Russian Academy of Sciences, Miklukho-Maklaya St. 16/10, 117997 GSP, Moscow B-437, Russia.

 $^{^{}m c}$ Moscow Politechnical University, Bolshaya Semenovskaya Str., 38, $\,$ 107023, Moscow, Russia.

[†] Corresponding author: mikle@irims.ru

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