

Synthesis of Novel Glycyrrhizic and Glycyrrhetic Acid Derivatives with Positive Antiviral Activity

Rimma M. Kondratenko¹, Lidia A. Baltina¹, Lilia R. Mikhailova¹, Lia A. Baltina¹, Olga V. Stolyarova¹, Genrikh A. Tolstikov¹, Sergey A. Nepogoev², Robert A. Field², Olaf Kunert³, Olga A. Plysunova⁴, Andrey G. Pokrovskiy⁴, Herold Hover⁵, and Jindrich Cinatl⁵

¹*Institute of Organic Chemistry Ufa Research Centre of RAS,
Prospekt Oktyabrya, 71, 450054, Ufa, Russia;
fax: +73472356066; e-mail: baltina@anrb.ru;*

²*School of Chemical Sciences and Pharmacy University of East Anglia, Norwich NR4 7TJ, UK;*

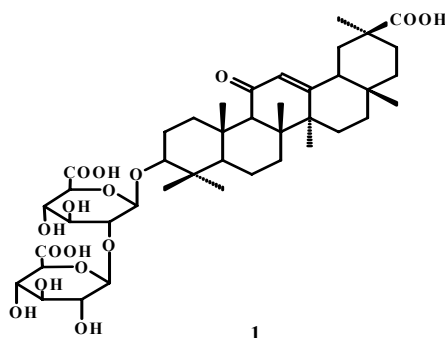
³*Institut für Pharmazeutische Chemie und Pharmazeutische Technologie,
Karl-Franzens-Universität Graz, Universitätsplatz 1, A-8010 Graz, Austria;*

⁴*State Scientific Centre "Vector", Novosibirsk;*

⁵*Institute of Medical Virology, Frankfurt University Medical School,
Paul-Enrich Str., 40, D-60596, Frankfurt, Germany*

Triterpenes and triterpene glycosides are natural compounds widely distributed in different plants. Combinatorial chemistry of high triterpenoids is a perspective route to design novel bioactive compounds for medicine.

To continue of our studies [1] in the field of transformations of Glycyrrhizic Acid (GL) (1) and its aglycon, Glycyrrhetic Acid (GLA) to be the main components of Glycyrrhiza Radix, we synthesised novel derivatives for the structure-antiviral activity studies. GL and GLA derivatives containing amino acids, hydrazides, heterocycles, D-glucosamine were produced by using simple protocols. Glycosylation of 18 β -GLA and its related compounds was studied by using iodine containing activators. 3-O- α - and - β -glycosides of 18 β - and 18 α -GLA, 11-desoxo- and 18,19-dehydro-derivatives to be modified GL analogues were produced. Chemical modifications of 11-desoxo-GL analogues were carried out by the introduction of bioactive esters, amides and aminoacids.



Transformations of ring A and oxidation of 11-desoxo-GLA in different conditions was studied. Among GL and GLA derivatives produced potent inhibitors of HIV-1 and SARS coV were found.

1. Baltina L.A. Chemical modification of Glycyrrhizic acid as a route to new bioactive compounds for medicine. *Current Med Chem* 2003; 10: 155-171.

This work was supported by the RFBR and Austria grant 03-03-20004 BNTS_a and grant 1488.2003.03 (Scientific School).