

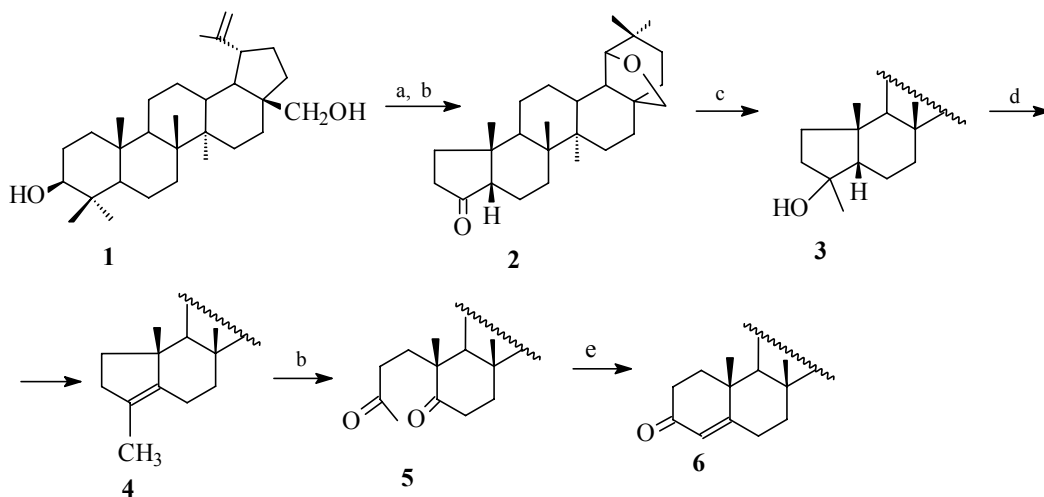
Synthesis of a New Triterpenoid – Steroid Hybrid Molecule from Betulin

N. I. Medvedeva, O. B. Flekhter, F. Z. Galin, G. A. Tolstikov

*Institute of Organic Chemistry, Ufa Center of the Russian Academy of Sciences,
71 Prospect Oktyabrya, 450054 Ufa, Russia
Fax(3472)356066
e-mail: obf@anrb.ru*

Earlier it was found, that some synthetic triterpenoids with modified ring A (for example 2-cyano-3,12-dioxolean-1,9-dien-28-oic acid, CDDO) induce differentiation and apoptosis of human leukemia cells [1]. Thus the search of new compounds among oleanane triterpenoids with potential biological activity is a very attractive area.

Here we report for the first time the synthesis of a new triterpenoid-steroid hybrid molecule from available lupane triterpenoid betulin. A-Nor-5 β H-19 β ,28-epoxy-18 α -olean-3-on **2**, received from betulin **1** via three steps, under treatment with MeMgI gave 3-hydroxy-3-methyl-derivative **3**. The following dehydration of **3** with POCl₃ in pyridine and ozonolysis of olefin **4** to diketone **5** were used. Aldol condensation of **5** in 5% NaOH/MeOH led to the new triterpenoid – steroid hybride molecule 23,24-dinor-3-oxo-4-en-19 β ,28-epoxy-18 α -olean **6**.



Reagents: a. 1) CF₃COOH, CH₂Cl₂; 2) PCl₅, 0⁰C, benzene/toluene; b. O₃, -60⁰C, CH₂Cl₂, Zn/AcOH;
c. MeMgI, ether; d. POCl₃/Py; e. NaOH/MeOH

[1] T. Ikeda, Y. Nakata, F. Kamura et al., *Mol. Cancer Ther.* **2004**, 3(1): 39-45.

The work was supported by the Russian Foundation for Basic Research (project no. 02-03-81007), grants from President of Russian Federation for supporting of young Russian scientists and leading scientific schools (project no. 543.2003.03, 1488.2003.3). OBF is grateful to the Science Support Foundation.