

Synthesis of 1'-N-homoazanucleosides as potential transition state analogues and antiviral agents

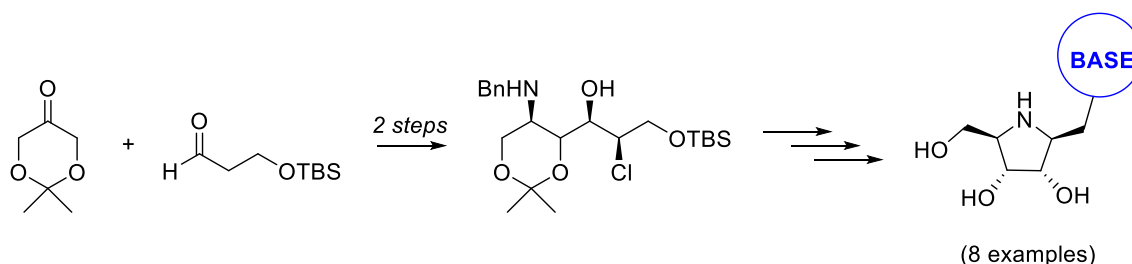
Jakob Bouton, Serge Van Calenbergh

Laboratory for Medicinal Chemistry (FFW), Ghent University, Ottergemsesteenweg 460,
9000 Ghent, Belgium

✉ Jakob.Bouton@ugent.be

Azanucleosides are a class of nucleoside analogues in which the furanose ring has been replaced by a functionalized pyrrolidine derivative. Most of the known azanucleosides are either C-nucleosides, or homonucleosides, in which a methylene linker is placed between the pyrrolidine and the nucleobase. A very promising class of aza-C-nucleosides are the Immucillins, which act as transition state analogue enzyme inhibitors of several nucleosidase & nucleoside phosphorylase enzymes.^[1] Some of these Immucillins also exhibit potent antiviral activity; the adenosine analogue BCX-4430 is currently being developed as a potential treatment for Ebola virus disease.^[2]

Inspired by the intriguing biological activities of known azanucleosides, a series of new 1'-N-homoazanucleosides was synthesized. These were constructed with a variety of nucleobases and nucleobase analogues and the synthesized compounds are currently being evaluated for antibacterial, antiprotozoal, antiviral and antitumoural activity.



References

- [1] Evans G. B., Schramm V. L., Tyler P. C., *Curr. Med. Chem.*, **2015**, 22, 3097
[2] Warren, T. K. *et al.*, *Nature*, **2014**, 508, 202