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Synthetic Study on ADP-ribosylation

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Proposition (Stellingen)

Accompanying the thesis

Synthetic study on ADP-ribosylation

1. The use of “parobiose” for the disaccharide in linear PAR and “parotriose” for the trisaccharide in branched PAR favors the scientific communication.
This Thesis
2. A solid-phase synthesis of ADPr-oligomers is preferred compared to a solution-phase approach.
J. Am. Chem. Soc. 2015, 137, 10, 3558, This thesis.
3. The branched structure elements in poly-ADP-ribose, of which the function is unknown, may determine the overall conformation of poly-ADP-ribose.
Miwa et al, J. Bio. Chem., 1981, 256, 2916.
4. The introduction of multiple pyrophosphate linkages is the biggest challenge in the synthesis of ADPr-oligomers.
This Thesis
5. Although it was discovered 40 years ago, branched poly-ADPr (PAR) has always been an enigma in the ADPr biology.
This Thesis
6. The detection of PAR (poly ADP-ribose), as a biomarker, could be useful for the future early diagnosis of neurodegeneration disorders.
Kam et al, Science, 2018, 362, 557.
7. Because the sensitivity of tumors for PARP inhibitors does not overlap with that for PARG inhibitors, a combination therapy is a viable option.
Pillay et al, Cancer Cell, 2019, 35, 519.
8. It is the duty of chemists to persuade ADPr-biologists to draw the structures of ADP-ribose with correct stereochemistry.

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