

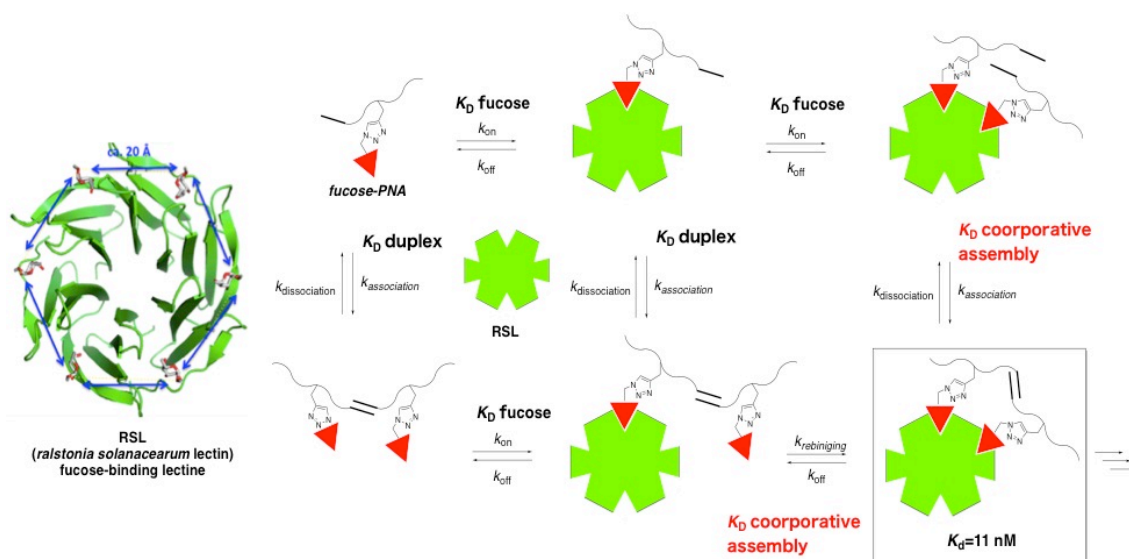
## Dynamic cooperative glycan assembly blocks binding of bacterial lectins to epithelial cells

T. Machida<sup>1</sup>, A. Novoa<sup>1</sup>, É. Gillon<sup>2</sup>, S. Zheng<sup>3</sup>, J. Claudinon<sup>3</sup>, T. Eierhoff<sup>3</sup>, A. Imberty<sup>2</sup>, W. Römer<sup>3</sup>, N. Winssinger<sup>1\*</sup>

<sup>1</sup>Department of Organic Chemistry, NCCR Chemical Biology, University of Geneva, 30 quai Ernest Ansermet, 1211 Geneva, Switzerland, <sup>2</sup>CERMAV UPR5301, CNRS, and Université Grenoble Alpes, BP 53, 38041 Grenoble cedex 9, France, <sup>3</sup>Faculty of Biology, Albert-Ludwigs-University Freiburg, Schänzlestraße 1, and Centre for Biological Signalling Studies (BIOSS), Albert-Ludwigs-University Freiburg, Schänzlestraße 18, 79104 Freiburg, G

Pathogenic bacterial infection to the host frequently utilizes lectin which recognizes glycan on cell surface of host. Lectin usually has multiple glycan-binding pockets and the multivalent inhibitor which simultaneously blocks multiple pockets is potent anti-bacterial medication strategy.

RSL was successfully blocked by conjugate with fucose and short peptide nucleic acid (PNA) with palindromic sequence ( $K_D=11$  nM) in which neither fucose nor PNA had comparable affinity (fucose:  $K_D=2200$  nM. PNA: GGCC, self hybridization  $K_D=3800$  nM). That suggested that host protein stabilize beneficial dimer formation. This conjugate had  $IC_{50}$  of 555 nM to inhibit the binding of fucose-binding lectin BambL to epithelial cells with efficiency of more than 700-fold compared to L-fucose.



1) T. Machida, A. Novoa, É. Gillon, S. Zheng, J. Claudinon, T. Eierhoff, A. Imberty, W. Römer, N. Winssinger, *Angew. Chem. Int. Ed.* **2017**, in press.