

# Synthesis of chemical tools to study the immune system

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### Propositions

## accompanying the thesis

### Synthesis of chemical tools to study the immune system

 The residual activity seemingly always displayed by caged TLR2/6 ligands cannot be reduced through modification of the polar surface area or molecular volume of the cage.

This thesis, Chapter 2

2. High-resolution structural data of receptor-ligand complexes are imperative for designing conditionally controlled ligands.

This thesis, Chapter 2 and 3

The ability to doubly functionalize a cage prospers its usability in a wide variety of experiments, even those for which a singly functionalized cage would seem to suffice.

This thesis, Chapter 2, 3, 4 and 5

4. Yields reported for large, insoluble, solid-phase derived molecules are often a reflection of unoptimized purification techniques rather than a reflection of the reaction efficiency.

This thesis, Chapter 2 and 5

- 5. The field of photon-mediated uncaging would benefit from a detailed description of the deprotection conditions and photon sources employed.
- The interdisciplinary nature of applying photo-protecting groups in a biological context implores scrutiny from all disciplines when performing root-causeanalysis of downstream hurdles.
- Uncaging conditions should, aside from being bio-orthogonal, also be virtually chemo-orthogonal, when the cage cannot be incorporated in the final step of caged ligand synthesis.

- 8. The activity of some TLR ligands is inversely correlated to their aqueous solubility.
- 9. The tradition to bind Ph.D. experiments in a single booklet with a forced overarching theme can result in the neglection of important data that happens to fall outside the theme.
- 10. The drive towards obtaining interesting biological data is often not parallel with the drive towards obtaining interesting synthetic routes.