## Characterization of Amyloid β aggregation via Supercritical Angle Fluorescence and Raman microscopy and spectroscopy

N. S. Münch<sup>1</sup>, S. Seeger<sup>1</sup>\*

<sup>1</sup>University of Zurich, Department of Chemistry, Winterthurerstrasse 190, 8057 Zürich

The two main forms of the Amyloid  $\beta$  (A $\beta$ ) peptide, A $\beta$ 1-40 and A $\beta$ 1-42 play an important role in Alzheimer's disease (AD), one of the main neurodegenerative diseases related to aging. The most important consequence of AD is the degradation of neural cells due to the aggregation of A $\beta$  into oligomers and fibrils with a subsequent accumulation outside the neurons [1]. Both A $\beta$  peptides play a different role in the AD process and in neurotoxicity. Therefore, a comparison of the aggregation of both peptides is crucial.

In this work, we studied the aggregation of A $\beta$  *in vitro*, directly on the lipid bilayer and compare the surface with the bulk solution, using Supercritical Angle Fluorescence (SAF) [2] and Raman (SAR) [3] microscopy and spectroscopy. The advantage of Raman over fluorescence is the labelfree method with the resolution of the peptide's secondary structure. The experiments clearly show A $\beta$ 1-42 aggregates stronger at the surface where more  $\alpha$ -helices are present, whereas A $\beta$ 1-40 forms mostly  $\beta$ -sheets. In addition, we studied the effect of calcium ions and confirmed a structural change of the peptides.

[1] Hossein Ashrafian, Elaheh Hadi Zadeh, Rizwan Hasan Khan, *International Journal of Biological Macromolecules*, **2021**, 167, 382-394.

[2] Doriel Verdes, Thomas Ruckstuhl, Stefan Seeger, *Journal of Biomedical Optics*, **2007**, 12(3), 034012.

[3] Diana Serrano, Stefan Seeger, *Light: Science & Applications*, **2017**, 6, e17066.