

# Parkinson's protein $\alpha$ -synuclein : membrane interactions and fibril structure

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## Citation

Kumar, P. (2017, June 27). Parkinson's protein  $\alpha$ -synuclein : membrane interactions and fibril structure. Casimir PhD Series. Retrieved from https://hdl.handle.net/1887/50076

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Author: Kumar, Pravin Title: Parkinson's protein  $\alpha$ -synuclein : membrane interactions and fibril structure Issue Date: 2017-06-27

# Stellingen

#### behorende bij het proefschrift

### Parkinson's Protein $\alpha$ -Synuclein: Membrane Interactions

#### and Fibril Structure

- 1. The conical shape of Cardiolipin and not its charge is the decisive factor for  $\alpha$ -Synuclein ( $\alpha$ S) binding to the inner mitochondrial membrane. [Chapter 2]
- 2. Neither a model membrane with a charge density  $\rho = 1$  nor one with a low charge density ( $\rho \le 0.3$ ) is suitable to investigate the effect of phosphorylation on  $\alpha$ S-membrane binding. [Chapter 3]
- 3. The biggest obstacle to study the intrinsic fold of  $\alpha$ S in fibrils is the polymorphism of the fibrils. [Chapter 4 and 5]
- 4. A single technique is not sufficient to obtain a realistic picture of the fibril-fold of  $\alpha$ S. [chapter 5]
- 5. The abundant occurrence in Lewy bodies of  $\alpha$ S phosphorylated at position S129 may be a result of phosphorylation of aggregated  $\alpha$ S at position S129 rather than phosphorylation of monomeric  $\alpha$ S. [Oueslati A.J. Parkinsons Dis. 2016;6:39-51, Paleologou K. et al. J. Neurochem. 2010;30:3184-3198]
- 6. The approach using lipid nanodiscs to study membraneprotein interactions described by Schuler et al. can be applied to investigate membrane fusion in vitro. [Schuler M.A. et al. Methods Mol. Biol. 2013;974:415-433]
- The sulfhydryl modification of the styrene-maleic acid copolymer (SMA-SH) described by Lindhoud et al. makes SMA-SH a potential target for spin labelling, which can be

used in the biophysical characterization of membrane proteins by EPR. [Lindhoud S. et al. Biomacromolecules 2016;17:1516-1522]

- 8. The O-GlcNAcylation (addition of a single monosaccharide Nacetyl-glucosamine) of  $\alpha$ S surprisingly inhibits the phosphorylation at position S129 but not at position S87. [Marotta N.P. et al. Nat.Chem. 2015;7:913-920]
- 9. Ignorance creates an illusion of confidence that knowledge does not.

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27-06-2017