



# 38<sup>a</sup> REUNIÃO ANUAL

Grupo de Estudos  
de Envelhecimento  
Cerebral e Demência

**17 e 18 de maio de 2024**

Escola de Medicina da Universidade do Minho  
Braga

**LIVRO DE RESUMOS**



Grupo De Estudos  
De Envelhecimento  
Cerebral e Demência

## CO-22 - BACTERIOPHAGES TO TACKLE ALZHEIMER'S DISEASE

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The first stages of Alzheimer's disease (AD) pathology involve the formation of toxic amyloid beta ( $A\beta$ ) oligomers, which can begin to form up to two decades before symptoms and other known disease markers appear. The ability to detect  $A\beta$  oligomers would allow early diagnosis, leading to improved disease management, care and prognosis through the use of therapeutic strategies targeting  $A\beta$  before irreparable brain damage occurs.

Peptides with affinity for  $A\beta$  may be the answer, as they are known to inhibit  $A\beta$  assembly and toxicity. However, peptides are highly unstable and do not cross the blood-brain barrier (BBB). To overcome this limitation, bacteriophages (phages) can be used as vectors for peptide delivery. Phages are viruses that selectively target and kill bacteria, safe for humans, easily modified to display a variety of biomolecules on their surface, and are known to be able to cross most biological barriers, including the BBB.

We have engineered phages to display  $A\beta$ -specific peptides and our results showed that the synthetic phages effectively identified  $A\beta$  aggregates in brain samples from AD-model mice from several ages, and AD patients, underlining their specificity towards oligomeric  $A\beta$ . We are validating the therapeutic potential of this phage-based tool in AD-mice, with promising exploratory results.

The outputs of this project will made available a theragnostic tool for early detection, monitorization and therapy of AD. This will have a positive impact on AD research and consequently on healthcare systems, society and economy, by reducing the overall costs and hurdles associated with this debilitating disease.