Oxygenation attenuates aggregation and neurotoxicity of amyloid beta (Aβ) peptide ~New Therapeutic Strategy for the Treatment of Alzheimer's Disease~

Summary;

Neurotoxicity of aggregated amyloid beta (A β) peptide is considered to be the pathogenesis of Alzheimer's disease. Although therapeutic approaches targeting A β have been intensively studied, the disease is currently incurable. Thus, a new approach should be developed to overcome the disease. A research group at The University of Tokyo designed and synthesized a new photocatalyst that can selectively oxygenate A β . The oxygenated A β exhibits remarkably low aggregation activity and neurotoxicity, indicating that photooxygenation of A β is able to attenuate A β toxicity. Improvement of such a photocatalyst toward biological application will lead to a new therapeutic strategy for the treatment of Alzheimer's disease.



<Figure 1> Concept of photooxygenation reaction for Aβ.

Aggregation of A β (blue) is inhibited by photocatalyzed oxygenation. A β is converted to oxygenated A β by binding with oxygen atoms (orange circles).



<Figure 2> Atomic force microscope (AFM) images of non-oxygenated (left) and oxygenated (right) A β . Fibrillar A β aggregates are observed when A β is not oxygenated (left), while they are not observed when A β is oxygenated (right). Thus, oxygenated A β is not aggregative.



<Figure 3> Attenuation of Aβ toxicity by photocatalyzed oxygenation.

Cell viability is decreased due to $A\beta$ toxicity when the neuronal cells are treated with non-oxygenated $A\beta$ (Compare a and b). On the other hand, cell viability recovered due to attenuated $A\beta$ toxicity when $A\beta$ is oxygenated by visible light irradiation in the presence of the photocatalyst (Compare b, c, and d).

Journal Information;

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(Program) JST ERATO(Project) Kanai Life Science Catalyst Project(Project Leader) Professor Motomu KANAI, The University of Tokyo

Contact Information;

Professor Motomu KANAI Graduate School of Pharmaceutical Sciences, The University of Tokyo 7-3-1 Hongo, Bunkyo-ku, Tokyo 113-0033, JAPAN e-mail kanai@mol.f.u-tokyo.ac.jp TEL +81-3-5841-4830 FAX +81-3-5684-5206