



ASX ANNOUNCEMENT

Actinogen presents academic poster at Australian Dementia Research Forum 2024 conference

Sydney, 3 June 2024. Actinogen Medical ASX:ACW (“ACW” or “the Company”) is pleased to announce that its Clinical Scientist Dr Jack Taylor is presenting an academic poster at the *Australian Dementia Research Forum* (ADRF2024) on the Gold Coast, Australia from 3 to 5 June 2024.

The theme for ADRF2024 is *Turning the Tide on Dementia*, with particular focus on exciting innovations in the field. The Forum is a meeting place for national and global experts to share the latest developments in dementia research, care and policy.

A copy of the poster is attached to this announcement.

The poster, entitled *Clinical pharmacology and development of Xanamem[®], a tissue specific inhibitor of 11 β -HSD1*, describes Xanamem’s clinical pharmacology, including Actinogen’s innovative approach to dose selection. The poster details how traditional pharmacokinetics and pharmacodynamics were paired with PET brain imaging and cognitive testing and other clinical data to confirm a target dose range of ≤ 10 mg daily.

Dr Dana Hilt, the Company's Chief Medical Officer said:

“Dr Taylor’s presentation summarizes the multiple streams of data supporting the selection of Xanamem doses of ≤ 10 mg for the treatment of cognitive impairment in a number of diseases. Via Inhibition of 11 β -HSD1, the enzyme synthesizing cortisol in brain, low doses of Xanamem potentially improve cognition and slow disease progression in patients with AD.

To our knowledge Xanamem is the first drug of this class to have compelling data showing robust central nervous system (CNS) target engagement, and positive procognitive effects in three independent, placebo-controlled trials.”

ENDS

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Announcement authorised by the Board of Directors of Actinogen Medical

About Actinogen Medical

Actinogen Medical (ACW) is an ASX-listed, biotechnology company developing a novel therapy for neurological and neuropsychiatric diseases associated with dysregulated brain cortisol. There is a strong association between cortisol and detrimental changes in the brain, affecting cognitive function, harm to brain cells and long-term cognitive health.

Cognitive function means how a person understands, remembers and thinks clearly. Cognitive functions include memory, attention, reasoning, awareness and decision-making.

Actinogen is currently developing its lead compound, Xanamem, as a promising new therapy for Alzheimer's Disease and Depression and hopes to study Fragile X Syndrome and other neurological and psychiatric diseases in the future. Reducing cortisol inside brain cells could have a positive impact in these and many other diseases. The cognitive dysfunction, behavioural abnormalities, and neuropsychological burden associated with these conditions is debilitating for patients, and there is a substantial unmet medical need for new and improved treatments.

Current Clinical Trials

The **XanaCIDD Phase 2a cognition & depression trial** is a double-blind, six-week proof-of-concept, placebo-controlled, parallel group design trial in 167 patients. Participants are evenly randomized to receive Xanamem 10 mg once daily or placebo, in some cases in addition to their existing antidepressant therapy, and effects on cognition and depression are assessed. Results are anticipated in early Q3 CY2024.

The **XanaMIA Phase 2b Alzheimer's disease trial** is a double-blind, 36-week treatment, placebo-controlled, parallel group design trial in 220 patients with mild to moderate AD and progressive disease, determined by clinical criteria and confirmed by an elevated level of the pTau181 protein biomarker in blood. Patients receive Xanamem 10 mg or placebo, once daily, and effects on cognition, function and progression of Alzheimer's disease are assessed. Thus, Xanamem is being assessed in this trial for its potential effects as a both a cognitive enhancer and a disease course modifier. Initial results from an interim analysis of the first 100 participants are anticipated in mid 2025.

About Xanamem

Xanamem's novel mechanism of action is to block the production of cortisol inside cells through the inhibition of the 11 β -HSD1 enzyme in the brain. Xanamem is designed to get into the brain after it is absorbed in the intestines upon swallowing.

Chronically elevated cortisol is associated with cognitive decline in Alzheimer's Disease and excess cortisol is known to be toxic to brain cells. Cognitive impairment is also a feature in Depression and many other diseases. Cortisol itself is also associated with depressive symptoms and when targeted via other mechanisms has shown some promise in prior clinical trials.

The Company has studied 11 β -HSD1 inhibition by Xanamem in more than 300 volunteers and patients, so far finding a statistically significant improvement in working memory and attention, compared with placebo, in healthy, older volunteers in two consecutive trials and clinically significant improvements in functional and cognitive ability in patients with biomarker-positive mild AD. Previously, high levels of target engagement in the brain with doses as low as 5 mg daily have been demonstrated in a human PET imaging study. A series of Phase 2 studies in multiple diseases is being conducted to further confirm and characterize Xanamem's therapeutic potential.

Xanamem is an investigational product and is not approved for use outside of a clinical trial by the FDA or by any global regulatory authority. Xanamem[®] is a trademark of Actinogen Medical.

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