ASX ANNOUNCEMENT

Professor Paul Maruff joins Actinogen CMO, Dr Dana Hilt, in a 'fireside chat' to review progress in evaluation of cognition, Alzheimer's disease and Xanamem's® clinical data

Prof Paul Maruff & Dr Dana Hilt neuroscience webinar: 11am Thursday 31 August 2023
Event registration: https://us02web.zoom.us/webinar/register/WN_TparOd53TUWkFUGeobwgTw

Sydney, 29 August 2023. Actinogen Medical ASX: ACW ("ACW" or "the Company") is pleased to announce that one of the world's leading neuroscience authorities and cognition experts, Professor Paul Maruff, will join Actinogen's Chief Medical Officer and neurologist Dr Dana Hilt MD to discuss recent progress in the Alzheimer's disease (AD) field and cognitive impairment associated with depressive disorder (CIDD) at **11am on Thursday 31 August 2023**.

This highly informative 'plain English' interview and discussion will focus on interpreting the various testing methods that have been applied to cognition in AD and CIDD and used to evaluate the efficacy of new drugs such as Xanamem.

Xanamem is ACW's promising novel once-a-day oral medication, currently under clinical development in a Phase 2a trial in patients with CIDD, and about to enter a new Phase 2b trial in patients with mild to moderate AD. It works on lowering brain cortisol and is one of only a few development programs that has demonstrated clinical activity in tests of cognition. Xanamem has positive data from three independent, controlled clinical trials to date.

Professor Paul Maruff is a Professor in Neuroscience at the Florey Institute of Neuroscience and in Psychology at Monash University, Melbourne Australia. He is also co-founder of Australian public company Cogstate Limited, which plays a major role in the assessment of Alzheimer's disease globally.

Professor Maruff has more than 30 years of experience in the assessment of cognition (how you think, remember and solve problems) and other tests used to assess AD and other diseases associated with cognitive impairment.

Dr Dana C. Hilt MD has more than 25 years of drug development experience, primarily of Central Nervous System (CNS) drugs. Dr Hilt has world-leading expertise and experience in Phases 1 to 4 of development for conditions including Alzheimer's disease, depression, Parkinson's disease and other neurologic and neuropsychiatric diseases.

Register now for this event on Thursday, 31 August at 11am:
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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 and Upcoming Clinical Trials

The **XanaCIDD Phase 2a depression trial** is a double-blind, six-week proof-of-concept, placebo-controlled, parallel group design trial in 160 patients. Patients 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.

The **XanaMIA Phase 2b Alzheimer's disease trial** is a double-blind, 36-week treatment, placebo-controlled, parallel group design trial in 330 patients with mild to moderate AD and progressive disease, determined by clinical criteria and confirmed by an elevated level of pTau181 protein in blood. Patients receive Xanamem 5 mg or 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.

About Xanamem

Xanamem's novel mechanism of action is to block the production of cortisol inside cells through the inhibition of the 11β-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 11β-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-

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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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