Proteinopathies such as Alzheimer's disease (AD) and Parkinson's disease (PD) are a group of disorders thought to be caused by abnormal folding or misfolding of beta amyloid (Ab) and α-synuclein, respectively. Their pathogenesis is not well understood due to unresolved molecular mechanisms. This is further complicated by the lack of proper natural disease models that might be effective in aiding the investigation of the molecular mechanisms underlying these disorders. Dogs spontaneously deposit human-type Ab as they age and thus are a natural higher mammalian model of aging. The canine Aβ precursor protein (APP) is virtually identical to human APP. Previous studies demonstrated that aging dogs spontaneously accumulate human-type Aβ and parallel declines in cognition. Further, the outcomes of immunotherapy studies in aged dogs has predicted human clinical trial outcomes; clearance of Aβ plaques with little cognitive benefits. In more recent work, we show that canine derived Aβ was toxic to human neuronal cell lines and led to aggregation of human Aβ. Eastern grey kangaroos (EGK) display a typical movement disorder presentation associated with grass phalaris poisoning. We show that this disorder, known as phalaris staggers displays a parkinsonian type syndrome with associated Parkinsoni-like signs and neuropathology, including synucleinopathy and neuromyopathy. Studies of proteinopathies have typically used transgenic mouse models and subsequently translated to human clinical trials. However, the success rate of these translational studies has been limited and unfortunately resulted in negative outcomes and some with adverse events. It is critical to identify and validate natural higher mammalian models of proteinopathies to investigate the molecular mechanisms underlying these disorders and test therapeutic outcomes prior to translation to human clinical trials.

Biography

Mourad Tayebi is an Associate Professor in Biomedical Sciences at the School of Medicine at Western Sydney University, Australia. He is an International Expert in the field of protein misfolding diseases, with specific focus on investigating the molecular mechanisms underlying pathogenic protein misfolding and characterizing the misfolding associated with these disorders. His team is very active in the development of early blood diagnostic test screen for Alzheimer's and effective therapies for neurodegenerative diseases.

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