

This invention is directed to novel aliphatic prolinamide derivatives of Formula I, and pharmaceutically acceptable salts, solvates, solvates of the salt and prodrugs thereof, useful in the prevention (e.g., delaying the onset of or reducing the risk of developing) and treatment (e.g., controlling, alleviating, or slowing the progression of) of age-related macular degeneration (AMD) and related diseases of the eye. These diseases include dry-AMD, wet-AMD, geographic atrophy, diabetic retinopathy, retinopathy of prematurity, polypoidal choroidal vasculopathy, and degeneration of retinal or photoreceptor cells. The invention disclosed herein is further directed to methods of prevention, slowing the progress of, and treatment of dry-AMD, wet-AMD, and geographic atrophy, diabetic retinopathy, retinopathy of prematurity, polypoidal choroidal vasculopathy, and degeneration of retinal or photoreceptor cells, comprising: administration of a therapeutically effective amount of compound of the invention. The compounds of the invention are inhibitors of HTRA1. Thus, the compounds of the invention are useful in the prevention and treatment of a wide range of diseases mediated (in whole or in part) by HTRA1. The compounds of the invention are also useful for inhibiting HTRA1 protease activity in an eye or locus of an arthritis or related condition.