

## **DNL758**

DNL758, a Peripheral Product discovered by Denali for which IND-enabling studies have been completed. DNL758 is a small molecule inhibitor of RIPK1 that does not penetrate the brain. Sanofi will lead clinical development activities for all systemic inflammatory diseases. The clinical trials are expected to begin in 2019.

The two lead molecules DNL747 and DNL758 target a critical signaling protein known as the receptor-interacting serine/threonine-protein kinase 1 (RIPK1) in the TNF receptor pathway, which regulates inflammation and cell death in tissues throughout the body. The companies plan to study DNL747 in Alzheimer's disease, amyotrophic lateral sclerosis and multiple sclerosis, and DNL758 in systemic inflammatory diseases such as rheumatoid arthritis and psoriasis.

Phase 1b and 2 clinical development costs for DNL747 will be fully funded by Sanofi for MS, ALS, and other neurological indications, except in Alzheimer's disease, which will be funded by Denali. Phase 3 trials for all neurological indications will be jointly funded by Sanofi (70%) and Denali (30%). Sanofi will fully fund the clinical development costs for DNL758 in systemic inflammatory diseases.