

# VX-765

Inflammatory Caspase-1 has a significant impact on AD-like pathophysiology and Caspase-1 inhibitor, VX-765, reverses cognitive deficits in AD mouse models. Here, a one-month pre-symptomatic treatment of Swedish/Indiana mutant amyloid precursor protein (APPsw/Ind) J20 and wild-type mice with VX-765 delays both APPsw/Ind- and age-induced episodic and spatial memory deficits. VX-765 delays inflammation without considerably affecting soluble and aggregated amyloid beta peptide (A $\beta$ ) levels. Episodic memory scores correlate negatively with microglial activation. These results suggest that Caspase-1-mediated inflammation occurs early in the disease and raise hope that VX-765, a previously Food and Drug Administration-approved drug for human CNS clinical trials, may be a useful drug to prevent the onset of cognitive deficits and brain inflammation in AD <sup>1)</sup>.

---

VX765 can counteract neurological damage after TBI by reducing pyroptosis and HMGB1/TLR4/NF- $\kappa$ B pathway activities. VX765 may have a good therapeutic effect on TBI <sup>2)</sup>

<sup>1)</sup>

Flores J, Noël A, Foveau B, Beauchet O, LeBlanc AC. Pre-symptomatic Caspase-1 inhibitor delays cognitive decline in a mouse model of Alzheimer disease and aging. Nat Commun. 2020 Sep 11;11(1):4571. doi: 10.1038/s41467-020-18405-9. Erratum in: Nat Commun. 2021 Apr 9;12(1):2271. PMID: 32917871; PMCID: PMC7486940.

<sup>2)</sup>

Sun Z, Nyanzu M, Yang S, Zhu X, Wang K, Ru J, Yu E, Zhang H, Wang Z, Shen J, Zhuge Q, Huang L. VX765 Attenuates Pyroptosis and HMGB1/TLR4/NF- $\kappa$ B Pathways to Improve Functional Outcomes in TBI Mice. Oxid Med Cell Longev. 2020 Apr 15;2020:7879629. doi: 10.1155/2020/7879629. PMID: 32377306; PMCID: PMC7181015.

From:

<https://neurosurgerywiki.com/wiki/> - **Neurosurgery Wiki**



Permanent link:

<https://neurosurgerywiki.com/wiki/doku.php?id=vx-765>

Last update: **2024/06/07 02:58**