

# Produktinformation



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Diagnostik & molekulare Diagnostik



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# Anti-Amyloid beta A4 protein [LY2062430 (Solanezumab, hu266)] Bulk Size Ab04185-10.3-BT

This antibody was created using our proprietary Fc Silent™ engineered Fc domain containing key point mutations that abrogate binding to Fc gamma receptors.

This is a reformatted human IgG1 Fc Silent Fc Silent™ antibody, based on the original human IgG1 format, created for improved compatibility with existing reagents, assays and techniques.

**Isotype and Format:** Human IgG1, Fc Silent<sup>™</sup>, Kappa Clone Number: LY2062430 (Solanezumab, hu266)

**Alternative Name(s) of Target:** APP; Aβ peptide; Abeta; A-beta; Aβ; Aβ42; amyloid beta peptide; Alzheimer disease amyloid protein; Beta-amyloid precursor protein; Protease nexin-II; humanized m266.2

**UniProt Accession Number of Target Protein:** P05067

Published Application(s): therapeutic, ELISA

Published Species Reactivity: Human

**Immunogen:** The parental mouse antibody 266 was generated by immunization of mice with a peptide composed of residues 13-28 of human  $A\beta$  peptide. The original humanized version of the antibody was generated by grafting CDRs of the mouse antibody onto human framework regions.

**Specificity:** This antibody recognizes amino acids 13-28 of amyloid beta and only recognized soluble form of the amyloid beta peptide. It functions as a cell surface receptor and performs physiological functions on the surface of neurons relevant to neurite growth, neuronal adhesion and axonogenesis. Interaction between APP molecules on neighboring cells promotes synaptogenesis.

**Application Notes:** The human IgG1 version of this antibody binds amyloid beta peptide Aβ1-42 with a binding affinity of Kd= 4pM in an in vitro BIAcore assay. The binding characterization of this antibody towards Aβ1-42-BSA conjugate was done using ELISA (US8591894). A study in patients with Alzheimer's disease (AD) suggested that a single dose of solanezumab was generally well tolerated, except that mild self-limited symptoms consistent with infusion reactions occurred in few patients when higher doses are given. A dose-dependent change in plasma and CSF Abeta was also observed (PMID: 20375655). The original mouse antibody 266 slowed Aβ accumulation in the brain but failed to deplete Aβ plaques in animal studies (PMID: 11438712). This humanized antibody is likely to have impeded the efflux of soluble Aβ from the brain in patients owing to the formation of Aβ-antibody complexes in the brain interstitial fluid and cerebrospinal fluid, as suggested from animal studies (PMID: 24638135). Phase 1 and 2 studies of

solanezumab revealed evidence of target engagement by dose-dependent increases in plasma and CSF total A $\beta$  (PMID: 20375655; 22672770). In the phase 2 study of mild to moderate AD, 12 weeks of solanezumab treatment yielded a dose-dependent increase in CSF-free A $\beta$ 42, suggesting a shift in equilibria sufficient to mobilize A $\beta$ 42 from plaques (PMID: 22672770). In the first phase III studies, solanezumab did not demonstrate significant benefit for the primary outcomes in either study but showed a favorable safety profile (PMID: 26238576).

**Antibody First Published in:** Siemers et al. Safety and changes in plasma and cerebrospinal fluid amyloid beta after a single administration of an amyloid beta monoclonal antibody in subjects with Alzheimer disease. Clin Neuropharmacol. 2010 Mar-Apr;33(2):67-73. PMID:20375655

**Note on publication:** This study evaluates the safety and tolerance of solanezumab after single administration in patients with Alzheimer's disease.

#### **Product Form**

**Size:** 1 mg Purified antibody in bulk size. **Purification:** Protein A affinity purified

**Supplied In:** PBS only.

**Storage Recommendation:** Store at 4°C for up to 3 months. Note, this antibody is provided without added preservatives, it is therefore recommed this antibody be handled under sterile conditions. For longer storage, aliquot and store at -20°C.

Concentration: 1 mg/ml.

Important note – This product is for research use only. It is not intended for use in therapeutic or diagnostic procedures for humans or animals.