Amitriptyline (hydrochloride) is supplied as a crystalline solid. A stock solution may be made by dissolving the amitriptyline (hydrochloride) in the solvent of choice. Amitriptyline (hydrochloride) is soluble in organic solvents such as ethanol, DMSO, and dimethyl formamide, which should be purged with an inert gas. The solubility of amitriptyline (hydrochloride) in these solvents is approximately 25 mg/ml.

Further dilutions of the stock solution into aqueous buffers or isotonic saline should be made prior to performing biological experiments. Ensure that the residual amount of organic solvent is insignificant, since organic solvents may have physiological effects at low concentrations. Organic solvent-free aqueous solutions of amitriptyline (hydrochloride) can be prepared by directly dissolving the crystalline solid in aqueous buffers. The solubility of amitriptyline (hydrochloride) in PBS, pH 7.2, is approximately 0.5 mg/ml. We do not recommend storing the aqueous solution for more than one day.

**Description**

Amitriptyline is a first generation tricyclic antidepressant.\(^1\) It inhibits serotonin (5-HT) uptake by isolated human platelets by 39 and 68% when used at concentrations of 1 and 4 µg/ml, respectively, and inhibits norepinephrine uptake in rat brain by 77% when administered at a dose of 10 mg/kg.\(^2,3\) It is also an antagonist of the 5-HT receptor subtype 5-HT\(_{2A}\) (K\(_i\) = 20 nM), as well as histamine H\(_1\), muscarinic acetylcholine, and \(\alpha_1\)-adrenergic receptors (K\(_i\)s = 1.1, 18, and 27 nM, respectively).\(^4-6\) Amitriptyline (5-25 µM) prevents acid sphingomyelinase activation and subsequent ceramide release induced by infection with replication-deficient vesicular stomatitis virus pseudoviral particles (pp-VSV) presenting the severe acute respiratory coronavirus 2 (SARS-CoV-2) spike protein in Vero cells, an effect that can be overcome with exogenous application of C16 ceramide (Item No. 10681) or recombinant human acid sphingomyelinase.\(^7\)

Formulations containing amitriptyline have been used in the treatment of depression and nerve pain. This product is also available as an analytical reference standard (Item Nos. 19029 | 19031).

**References**