

Catalog # 10-1439 ML-SA1

CAS# 332382-54-4

 $\hbox{$2$-[2-(3,4-Dihydro-2,2,4-trimethyl-1(2H)-quinolinyl)-2-oxoethyl]-1H-isoindole-1,3(2H)-dione} \\ Lot \# X105330$

TRPML 1,2 and 3 channel activator. Does not activate TRPM2, TRPV2, TRPV3, TRPC6 or TRPA1 channels. ML-SA1 robustly activates whole cell TRPML1-4A and whole-endolysosome TRPML1 and is comparable (10 μ M) to the effect of the endogenous TRPML agonist, PI(3,5)P₂ (1 μ M). Thus it is a useful chemical tool for studying the functions of TRPMLs. It induces TRPML-mediated Ca²⁺ release from lysosomes which corrects trafficking defects and reduces cholesterol accumulation in Niemann-Pick type C macrophages. Reduces intralysosomal Ca²⁺ level rescuing abnormal lysosomal storage in FIG4-deficient cells.

- 1) Grimm et al. (2010), Small molecule activators of TRPML3; Mol. Chem. Biol., 17 135
- 2) Shen et al. (2012), Lipid storage disorders block lysosomal trafficking by inhibiting a TRP channel and lysosomal calcium release; Nature Commun., **3** 731
- 3) Weiss et al. (2012), Cross-talk between TRPML1 channel, lipids and lysosomal storage diseases; Commun. Integer. Biol., 5 111
- 4) Zou et al. (2015), Reactivation of lysosomal Ca2+ Efflux Rescues Abnormal Lysosomal Storage in FIG4-Deficient Cells; J. Neurosci.. **35** 6801

PHYSICAL DATA

Molecular Weight: 362.43

Molecular Formula: C₂₂H₂₂N₂O₃

Purity: 98% by HPLC

NMR: (Conforms)

DMSO (up to 20 mg/ml with warming) or Ethanol (up to 10 mg/ml)

Physical Description: Tan solid

Solubility:

Storage and Stability: Store as supplied at room temperature for up to 1 year from the date of purchase. Solutions in

DMSO or ethanol may be stored at -20°C for up to 3 months.

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