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Produktinformation

- Forschungsprodukte & Biochemikalien
- Zellkultur & Verbrauchsmaterial
- Diagnostik & molekulare Diagnostik
- Laborgeräte & Service

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Lieferung & Zahlungsart

siehe unsere Liefer- und Versandbedingungen

Zuschläge

- Mindermengenzuschlag
- Trockeneiszuschlag
- Gefahrgutzuschlag
- Expressversand

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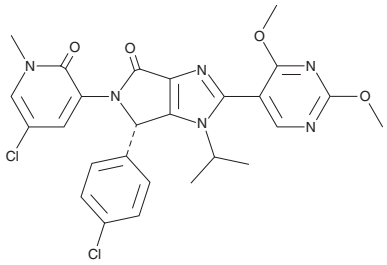
PRODUCT INFORMATION

Siremadlin

Item No. 38516



CAS Registry No.:	1448867-41-1
Formal Name:	(6S)-5-(5-chloro-1,2-dihydro-1-methyl-2-oxo-3-pyridinyl)-6-(4-chlorophenyl)-2-(2,4-dimethoxy-5-pyrimidinyl)-5,6-dihydro-1-(1-methylethyl)-pyrrolo-[3,4-d]imidazol-4(1H)-one
Synonyms:	HDM201, NVP-HDM201
MF:	$C_{26}H_{24}Cl_2N_6O_4$
FW:	555.4
Purity:	≥98%
UV/Vis.:	λ_{max}: 223, 336 nm
Supplied as:	A solid
Storage:	-20°C
Stability:	≥4 years

Information represents the product specifications. Batch specific analytical results are provided on each certificate of analysis.

Laboratory Procedures

Siremadlin is supplied as a solid. A stock solution may be made by dissolving the siremadlin in the solvent of choice, which should be purged with an inert gas. Siremadlin is soluble in organic solvents such as ethanol, DMSO, and dimethyl formamide. The solubility of siremadlin in these solvents is approximately 15, 10, and 20 mg/ml, respectively.

Description

Siremadlin is an inhibitor of the protein-protein interaction between the E3 ubiquitin ligase MDM2 and p53 (IC_{50} = 0.21 nM).[1] It selectively inhibits the MDM2-p53 protein-protein interaction over MDM4-p53, YAP1-TEAD4, PCSK9-LDLR, and Bcl-2-Bak interactions in time-resolved FRET (TR-FRET) assays (IC_{50}s = 3.3, >150, >50, and >50 µM, respectively). Siremadlin inhibits the growth of SJSA-1 osteosarcoma cells (GI_{50} = 38 nM). It also induces apoptosis in SJSA-1 cells when used at a concentration of 100 nM. *In vivo*, siremadlin (27 mg/kg) induces tumor regression and prevents tumor regrowth in an SJSA-1 rat xenograft model and an HSAX2655 patient-derived xenograft (PDX) rat liposarcoma model.

Reference

1. Jeay, S., Ferretti, S., Holzer, P., *et al.* Dose and schedule determine distinct molecular mechanisms underlying the efficacy of the p53-MDM2 inhibitor HDM201. *Cancer Res.* **78(21)**, 6257-6267 (2018).

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