

# **Bioactive Molecules, Building Blocks, Intermediates**

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# **Data Sheet**

Product Name:	CC-885
Cat. No.:	CS-0021543
CAS No.:	1010100-07-8
Molecular Formula:	C22H21CIN4O4
Molecular Weight:	440.88
Target:	Ligand for E3 Ligase
Pathway:	PROTAC
Solubility:	DMSO : 67.5 mg/mL (153.10 mM; Need ultrasonic)

# **BIOLOGICAL ACTIVITY:**

CC-885 is a cereblon (**CRBN**) modulator with potent anti-tumour activity. IC50 & Target: CRBN<sup>[1]</sup>. **In Vitro**: Acute myeloblatlic leukemia (AML) cell lines, human liver epithelial cell line (THLE-2) and human peripheral blood mononuclear cells (PBMC) are treated with varying concentrations of CC-885, with IC<sub>50</sub>s of  $10 \times ^{-6}$ -1  $\mu$ M. The effect of CC-885 on cell proliferation in AML cell lines, THLE-2 and human PBMC is more powerful than Lenalidomide and Pomalidomide with IC<sub>50</sub>s>10  $\mu$ M. To address whether the cereblondependent degradation of GSPT1 is responsible for the cytotoxic effects of CC-885, a GSPT1 mutant that retains its normal function, but loses CC-885-dependent cereblon binding, is used to distinguish the role of GSPT1 from that of other substrates. CC-885 is tested in 293T HEK cells stably expressing the CC-885-sensitive or -resistant GSPT1 variants. Overexpression of a resistant variant GSPT1  $\Delta$ (1–138)/(G575N) completely abrogate the CC-885-induced anti-proliferation, whereas overexpression of a CC-885-sensitive variant GSPT1 $\Delta$ (1-138) only confer partial protection. Similar results are obtained in AML cell lines<sup>[1]</sup>.

### PROTOCOL (Extracted from published papers and Only for reference)

**Cell Assay:** <sup>[1]</sup>Human cancer cell lines cultured in the growth medium are seeded into black 384-well plates containing DMSO or test compounds such as **CC-885** ( $10 \times ^{-6}$ -1  $\mu$ M). The seeding density for each cell line is optimized to allow the cell growth in the linear range during a 3-day culture period. To test the compound effect on cell proliferation in acute myeloid leukaemia (AML) cell lines, 5,000 to 10,000 cells per well in 200  $\mu$ l complete culture media are seeded into black 96-well plates containing DMSO or test compounds such as CC-885. After 48 or 72 h, cell proliferation is assessed using the CellTiter-Glo (CTG) Luminescent Cell Viability Assay<sup>[1]</sup>.

### **References:**

[1]. Mary E. Matyskiela, et al. A novel cereblon modulator recruits GSPT1 to the CRL4CRBN ubiquitin ligase. Nature. 2016 Jul 14;535(7611):252-7.

### **CAIndexNames:**

Urea, N-(3-chloro-4-methylphenyl)-N'-[[2-(2,6-dioxo-3-piperidinyl)-2,3-dihydro-1-oxo-1H-isoindol-5-yl]methyl]-

# SMILES:

O = C(NCC1 = CC2 = C(C(N(C(CC3)C(NC3 = O) = O)C2) = O)C = C1)NC4 = CC = C(C)C(CI) = C4

Caution: Product has not been fully validated for medical applications. For research use only.

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