Proteins

# **BL-918**

Cat. No.: HY-124729 CAS No.: 2101517-69-3 Molecular Formula:  $C_{23}H_{15}F_{8}N_{3}OS$ Molecular Weight: 533.44

Target: ULK; Autophagy Pathway: Autophagy

Storage: Powder -20°C 3 years

2 years

-80°C In solvent 2 years

> -20°C 1 year

**Product** Data Sheet

## **SOLVENT & SOLUBILITY**

In Vitro DMSO:  $\geq 250 \text{ mg/mL} (468.66 \text{ mM})$ 

\* "≥" means soluble, but saturation unknown.

Preparing Stock Solutions	Solvent Mass Concentration	1 mg	5 mg	10 mg
	1 mM	1.8746 mL	9.3731 mL	18.7463 mL
	5 mM	0.3749 mL	1.8746 mL	3.7492 mL
	10 mM	0.1875 mL	0.9373 mL	1.8746 mL

Please refer to the solubility information to select the appropriate solvent.

In Vivo

1. Add each solvent one by one: 10% DMSO >> 90% corn oil Solubility: ≥ 2.08 mg/mL (3.90 mM); Clear solution

# **BIOLOGICAL ACTIVITY**

Description BL-918 is an orally active UNC-51-like kinase 1 (ULK1) activator with an EC $_{50}$  of 24.14 nM. BL-918 exerts its cytoprotective autophagic effect by targeting ULK complex. BL-918 has the potential for Parkinson's disease (PD) treatment<sup>[1]</sup>.

IC<sub>50</sub> & Target ULK1 ULK1

> 24.14 nM (EC50) 0.719 μM (Kd)

In Vitro BL-918 (compound 33i) binds to ULK1 with a high binding affinity  $(K_D=0.719 \,\mu\text{M})^{[1]}$ .

BL-918 (5 μM; for 24 hours) induces autophagy in Neuron-Like SH-SY5Y cells<sup>[1]</sup>.

BL-918 (0.5-50 μM; for 24 hours) can partially reverse MPP+-induced cell death, which is determined by enhancing cell viability<sup>[1]</sup>.

BL-918 (5 μM; for 6-36 hours) time-dependently elevates the expression levels of LC3-II, Beclin-1, and its phosphorylation

status, whereas reduces the level of the selective autophagy substrate SQSTM1/p62. BL-918 elevates Ser317 and Ser555 phosphorylation of ULK1, as well as decreases Ser757 phosphorylation of ULK1<sup>[1]</sup>.

MCE has not independently confirmed the accuracy of these methods. They are for reference only.

## Cell Autophagy Assay<sup>[1]</sup>

cett/tatophagy/tosay		
Cell Line:	SH-SY5Y cells	
Concentration:	5 μΜ	
Incubation Time:	For 24 hours	
Result:	Induced Autophagy.	
Cell Viability Assay <sup>[1]</sup>		
Cell Line:	SH-SY5Y cells	
Concentration:	0.5, 5, 50 μΜ	
Incubation Time:	For 24 hours	
Result:	Could partially reverse MPP <sup>+</sup> -induced cell death, which was determined by enhancing cell viability.	
Western Blot Analysis <sup>[1]</sup>		
Cell Line:	SH-SY5Y cells	
Concentration:	5 μΜ	
Incubation Time:	6, 12, 24, 36 hours	
Result:	Time-dependently elevated the expression levels of LC3-II (a key marker of autophagy), Beclin-1, and its phosphorylation status, whereas reduced the level of the selective autophagy substrate SQSTM1/p62.	

#### In Vivo

BL-918 (compound 33i; 20, 40, or 80 mg/kg/day; oral gavage; began 2 days before the first injection of saline/MPTP and continuously maintained for 5 days after the last injection of saline/MPTP) attenuates the loss of DA and its metabolites. BL-918 obviously decreases the levels of dopamine (DA), 3,4-dihydroxyphenylacetic acid (DOPAC), and homovanillic acid (HVA) [1].

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Animal Model:	Male C57BL/6 mice (eight-week old) weighing between 20 and 25 $\mathrm{g}^{[1]}$	
Dosage:	20, 40, or 80 mg/kg	
Administration:	Oral gavage; daily; began 2 days before the first injection of saline/MPTP and continuously maintained for 5 days after the last injection of saline/MPTP	
Result:	Attenuated the loss of DA and its metabolites.	

### **REFERENCES**

[1]. Ouyang L, et al. Small-Molecule Activator of UNC-51-Like Kinase 1 (ULK1) That Induces Cytoprotective Autophagy for Parkinson's Disease Treatment. J Med Chem. 2018 Apr 12;61(7):2776-2792.

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 $\label{lem:caution:Product} \textbf{Caution: Product has not been fully validated for medical applications. For research use only.}$ 

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