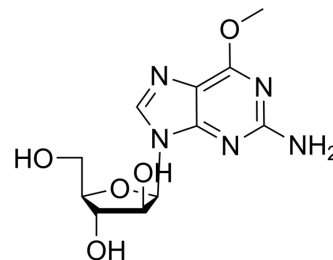


## Nelarabine

<b>Cat. No.:</b>	HY-13701		
<b>CAS No.:</b>	121032-29-9		
<b>Molecular Formula:</b>	C <sub>11</sub> H <sub>15</sub> N <sub>5</sub> O <sub>5</sub>		
<b>Molecular Weight:</b>	297.27		
<b>Target:</b>	Nucleoside Antimetabolite/Analog; Apoptosis		
<b>Pathway:</b>	Cell Cycle/DNA Damage; Apoptosis		
<b>Storage:</b>	Powder	-20°C	3 years
		4°C	2 years
	In solvent	-80°C	2 years
		-20°C	1 year



### SOLVENT & SOLUBILITY

#### In Vitro

DMSO : 100 mg/mL (336.39 mM; Need ultrasonic)  
 H<sub>2</sub>O : 10 mg/mL (33.64 mM; Need ultrasonic)

Preparing Stock Solutions	Solvent		1 mg	5 mg	10 mg
	Concentration	Mass			
	1 mM		3.3639 mL	16.8197 mL	33.6395 mL
	5 mM		0.6728 mL	3.3639 mL	6.7279 mL
	10 mM		0.3364 mL	1.6820 mL	3.3639 mL

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

#### In Vivo

- Add each solvent one by one: PBS  
Solubility: 5 mg/mL (16.82 mM); Clear solution; Need ultrasonic and warming and heat to 60°C
- Add each solvent one by one: 10% DMSO >> 40% PEG300 >> 5% Tween-80 >> 45% saline  
Solubility: ≥ 2.5 mg/mL (8.41 mM); Clear solution
- Add each solvent one by one: 10% DMSO >> 90% (20% SBE-β-CD in saline)  
Solubility: ≥ 2.5 mg/mL (8.41 mM); Clear solution
- Add each solvent one by one: 10% DMSO >> 90% corn oil  
Solubility: ≥ 2.5 mg/mL (8.41 mM); Clear solution

### BIOLOGICAL ACTIVITY

#### Description

Nelarabine (506U78) is a nucleoside analogue and can be used for the research of T cell acute lymphoblastic leukemia (T-ALL)<sup>[1]</sup>.

#### In Vitro

Nelarabine (506U78) (0-20 μM; 48 h) induces cytotoxic effects in T-ALL cell lines<sup>[1]</sup>.

Nelarabine (5 or 2  $\mu\text{M}$ ; 48 h) promotes apoptosis in sensitive T-ALL cell lines and modulates PI3K/AKT/mTOR and MEK signaling<sup>[1]</sup>.

Nelarabine (10  $\mu\text{M}$ ; 0-48 h) resistance does not depend on expression of ENT1/2 transporters and is partly due to upregulation of PI3K, MEK, and Bcl2 signaling<sup>[1]</sup>.

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

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

Cell Line:	T-ALL cell lines
Concentration:	0-20 $\mu\text{M}$
Incubation Time:	48 h
Result:	Cell viability decreased in a concentration-dependent fashion, and the IC <sub>50</sub> values ranged between 2 and 5.5 $\mu\text{M}$ for sensitive cell lines (MOLT-4, HSB-2, P12, DND41, JURKAT).

#### Apoptosis Analysis<sup>[1]</sup>

Cell Line:	MOLT-4, JURKAT, P12-ICHIKAWA and DND41
Concentration:	5 $\mu\text{M}$ (2 $\mu\text{M}$ for MOLT-4 cells)
Incubation Time:	48 h
Result:	Detected a marked increase in the percentage of early apoptotic and/or late apoptotic cells.

#### Western Blot Analysis<sup>[1]</sup>

Cell Line:	MOLT-4, JURKAT, P12-ICHIKAWA and DND41
Concentration:	5 $\mu\text{M}$ (2 $\mu\text{M}$ for MOLT-4 cells)
Incubation Time:	0, 6, 16, 24 and 48 h
Result:	Documented a time-dependent cleavage of caspase 8, caspase 9, caspase 3, and poly(ADP-ribose) polymerase (PARP) in response to drug treatment. Induced a marked decrease of phosphorylated AKT at Ser473, S6 ribosomal protein (S6RP) at Ser235/236, and GSK3 $\beta$ at Ser9.

#### In Vivo

Nelarabine (506U78) (130 mg/kg/day; i.v.; 5 days) reduces leukemic burden and extends mouse survival in NSG mice xenografted with luciferase-expressing U937 cells<sup>[2]</sup>.

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

Animal Model:	NSG mice xenografted with luciferase-expressing U937 cells <sup>[2]</sup>
Dosage:	130 mg/kg/day
Administration:	Intravenous injection, 5 days
Result:	Reduced leukemic burden and extended mouse survival.

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- J Mol Med (Berl). 2019 Aug;97(8):1183-1193.

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## REFERENCES

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[1]. Lonetti A, et al. Improving nelarabine efficacy in T cell acute lymphoblastic leukemia by targeting aberrant PI3K/AKT/mTOR signaling pathway. J Hematol Oncol. 2016 Oct 24;9(1):114.

[2]. Wang H, et al. Repurposing Nelarabine to Induce Differentiation of Acute Myeloid Leukemia. Blood, 2020, 136: 26.

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

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