

Catalog # 10-4479 Mitapivat

CAS# 1260075-17-9 N-[4-[4-(Cyclopropylmethyl)piperazine-1-carbonyl]phenyl]quinoline-8-sulfonamide; AG-348 Lot # FBA8062



Mitapivat is a potent (AC₅₀ = 29 nM PKR; 37 nM PKM2; 33 nM PKL) allosteric activator of pyruvate kinase.¹ In clinical use for the treatment of hemolytic anemia² and a potential agent for the treatment for ß-thalassemia³ and sickle cell anemia⁴. Pyruvate kinase M2 regulates the rate-limiting step of glycolysis that shifts glucose metabolism from normal respiratory chain to lactate production in tumor cells and is, thus, an important target in cancer cell metabolism.⁵

- 1) Kung et al. (2017), AG-348 enhances pyruvate kinase activity in red blood cells from patients with pyruvate kinase deficiency; Blood, **130** 1347
- 2) Rab et al. (2021), AG-348 (mitapivat), an allosteric activator of red blood cell pyruvate kinase, increases enzymatic activity, protein stability, and adenosine triphosphate levels over a broad range of PKLR genotypes; Haematologica, **106** 238
- 3) Matte et al. (2021), The pyruvate kinase activator mitapivat reduces hemolysis and improves anemia in a ß-thalassemia mouse model; J. Clin. Invest., **131** e144206
- 4) Rab et al. (2021), Decreased activity and stability of pyruvate kinase in sickle cell disease: a novel target for mitapivat therapy; Blood, **137** 2997
- 5) Zahra et al. (2020), Pyruvate Kinase M2 and Cancer: The Role of PKM2 in Promoting Tumorigenesis; Front. Oncol. 10 159

PHYSICAL DATA

Molecular Weight:	450.56
Molecular Formula:	$C_{24}H_{26}N_4O_3S$
Purity:	99% by HPLC
	NMR: (Conforms)
Solubility:	DMSO (>50 mg/mL)
Physical Description:	Beige solid
Storage and Stability:	Store as supplied desiccated at -20°C for up to 2 years from the date of purchase.
	Solutions in DMSO may be stored at -20°C for up to 3 months.

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