

AhR agonist

Catalog codes: tlrl-kyn, tlrl-kyn-5 https://www.invivogen.com/kynurenine

For research use only

Version 21K03-MM

PRODUCT INFORMATION

Contents

L-Kynurenine is available in two pack sizes:

• tlrl-kyn: 1 x 10 mg • tlrl-kyn-5: 5 x 10 mg

Storage and stability

- L-Kynurenine is shipped at room temperature. Store at 15-25 °C.
- Upon resuspension, store at 15-25°C. Resuspended product is stable for 6 months when properly stored. DO NOT FREEZE.
- We recommend to protect this product from light.

Quality control

- Purity ≥ 95% (UHPLC)
- The absence of bacterial contamination (e.g. lipoproteins and endotoxins) has been confirmed using HEK-Blue™ TLR2 and HEK-Blue[™] TLR4 cells.
- The biological activity has been validated using cellular assays.

BACKGROUND

L-Kynurenine (β -Anthraniloyl-L-Alanine) is an endogenous agonist for the cytosolic aryl hydrocarborn receptor (AhR)¹. L-Kynurenine results from the catabolic conversion of tryptophan (Trp) by two enzymes, the indoleamine-2,3-dioxygenase (IDO1) and the tryptophan-2,3dioxygenase (TDO2). L-Kynurenine is the first byproduct of Trp metabolism generated via the enzymatic "kynurenine pathway", which produces other kynunerine derivatives². L-Kynurenine is implicated in the production of TGF-β and expansion of regulatory T cells through the AhR-Src-IDO1 pathway¹. An increase in the kynurenine/Trp ratio in tumors has been shown to correlate with cancer progression, corroborating immuno-suppressive functions for this AhR agonist². Yet, L-Kynurenine's mode of action on AhR is still unclear. Indeed, L-Kynurenine could be a low-affinity AhR pro-ligand that is slowly converted in high-affinity compounds acting as AhR agonists at subnanomolar concentrations. This conversion occurs independently of enzymes, when fresh crystalline L-Kynurenine is resuspended as a solution, and gives rise to trace-amounts of derivatives named TEACOPs (trace-extended aromatic condensation products)3.

1. Bessede A. et al., 2014. Aryl hydrocarbon receptor control of a disease tolerance defence pathway. Nature. 511:184-90. 2. Hubbard T.D. et al., 2015. Indole and tryptophan metabolism: endogenous and dietary routes to Ah receptor activation. Drug Metab. Dispos. 43:1522-35. 3. Seok, S.H. et al., 2018. Trace derivatives of kynurenine potently activate the aryl hydrocarbon receptor (AHR). J. Biol. Chem. 293:1994-2005.

PRODUCT DESCRIPTION

As stated above, L-Kynurenine may be the pro-ligand for AhR that is converted into trace-amounts derivatives with high-affinity for AhR. To avoid weighing and ensure the most accurate manipulation of this product, L-Kynurenine is available as 10 mg units. Each lot of L-Kynurenine is functionally tested in cellular assays using our HepG2-Lucia[™] AhR and HT29-Lucia[™] AhR reporter cells.

CHEMICAL PROPERTIES

CAS number: 2922-83-0

Synonyms: β-Anthraniloyl-L-Alanine, L-2-Amino-4-(2-aminophenyl)-

4-oxobutanoic acid

Solubility: 20 mg/ml (96 mM) in DMSO

Formula: C₁₀H₁₂N₂O₃

Molecular weight: 208.22 g/mol Structure:



METHODS

Preparation of 5 mg/ml (24 mM) stock solution

- 1. Add 1 ml of DMSO to 10 mg of L-Kynurenine. Mix by vortexing and transfer into a new tube.
- 2. Rinse the commercial vial with 1 ml of DMSO and add to the previously resuspended product. Mix by vortexing. The final volume in the new tube is 2 ml.
- 3. Use immediately or store at 15-25 °C until required. **DO NOT FREEZE.** Note: L-Kynurenine's potency is very low when just resuspended. Potency increases days post-resuspension.

Working concentration range: 10 µg/ml (48 µM) to 100 µg/ml (480 µM) for cell culture assays

AhR activation assav

Described below is a protocol to study AhR activation in HepG2-Lucia™ AhR cells which derive from the human HepG2 hepatoma cell line. These cells report AhR activation through the monitoring of human Cyp1a1-induced Lucia luciferase activity. For more information, visit https://www.invivogen.com/hepg2-lucia-ahr.

- 1. Prepare a 5-fold dilution of L-Kynurenine stock solution in DMSO to obtain a 1 mg/ml work solution.
- 2. Prepare further dilutions using sterile endotoxin-free water.
- 3. Add 20 µl of L-Kynurenine at 33 µg/ml (final concentration) per well of a flat-bottom 96-well plate.

Note: we recommend to include a control dilution of DMSO in your assay.

- 4. Add 180 µl of cell suspension (~20,000 cells) per well.
- 5. Incubate the plate at 37 °C in a 5% CO₂ incubator for 18-24 hours.
- 6. Monitor Lucia luciferase reporter protein production using a luciferase detection reagent, such as QUANTI-Luc™.

RELATED PRODUCTS

Product	Description (Cat. Code
CH-223191 FICZ HepG2-Lucia™ AhR Cells HT29-Lucia™ AhR Cells ITE QUANTI-Luc™	AhR inhibitor AhR ligand AhR hepatoma reporter cells AhR colon cancer reporter cell AhR ligand Lucia detection medium	



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