

## Catalog # 10-4699

## **Firsocostat**

CAS# 1434635-54-7

2-[1-[(2R)-2-(2-Methoxyphenyl)-2-9oxan-4-yloxy)ethyl]-5-methyl-6-(1,3-oxazol-2-yl)-2,4-dioxothieno[2,3-d]pyrimidin-3-yl]2-methylpropanoic acid; ND-630; NDI-010976; GS-0976
Lot # FBS2213

Firsocostat is a potent ( $IC_{50} = 1.7$  nM ACC1; 2.6 nM ACC2) allosteric protein-protein interaction inhibitor of acetyl-CoA carboxylase (ACC). It interacts with the ACC phosphopeptide acceptor and dimerization site to inhibit enzymatic activity resulting in decreased fatty acid synthesis and stimulation of fatty acid oxidation. It reduced hepatic steatosis, improves insulin sensitivity, reduces weight gain, and favorably affects dyslipidemia in rats. Firsocostat decreased hepatic steatosis and fibrosis markers in patients with nonalcoholic fatty liver disease. Directly impairs profibrinogenic activity of hepatic stellate cells (HSC) via

1) Harriman et al. (2016), Acetyl-CoA carboxylase inhibition by ND-630 reduces hepatic steatosis, improves insulin sensitivity, and modulates dyslipidemia in rats; Proc. Nat. Acad. Sci. USA, **113** E1796

prevention of induction of glycolysis and oxidative phosphorylation during HSC activation.<sup>3</sup>

- 2) Looma et al. (2018), GS-0976 Reduces Hepatic Steatosis and Fibrosis Markers in Patients With Nonalcoholic Fatty Liver Disease; Gastroenterology, **155** 1463
- 3) Bates et al. (2020), Acetyl-CoA carboxylase inhibition disrupts metabolic reprogramming during hepatic stellate cell activation; J. Hepatol., **73** 896

## PHYSICAL DATA

Molecular Weight: 569.63

Solubility:

Molecular Formula: C<sub>28</sub>H<sub>31</sub>N<sub>3</sub>O<sub>8</sub>S Purity: 98% by HPLC NMR: (Conforms)

DMSO (>25 mg/ml)

Physical Description: White 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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