

## NN1177

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|-----------------------------|---|
| <b>Cat. No.:</b>            | HY-P10032   |
| <b>Molecular Formula:</b>   | C <sub>206</sub> H <sub>323</sub> N <sub>51</sub> O <sub>66</sub>   |
| <b>Molecular Weight:</b>    | 4570.07   |
| <b>Sequence:</b>            | His-{Aib}-Gln-Gly-Thr-Phe-Thr-Ser-Asp-Leu-Ser-Lys-Tyr-Leu-Glu-Ser-Lys-Arg-Ala-Arg-Glu-Phe-Val-Gln-Trp-Leu-Leu-{Lys(gGlu-gGlu-Ser-Glu-Ser-gGlu-gGlu-C18 diacid)}-Thr-NH <sub>2</sub> |
| <b>Sequence Shortening:</b> | H-{Aib}-QGTFTSDLSKYLESKRAREFVQWLL-{Lys(γGlu-γGlu-Ser-Glu-Ser-γGlu-γGlu-C18 diacid)}-T-NH <sub>2</sub>   |
| <b>Target:</b>              | GCCR; GLP Receptor; Cytochrome P450   |
| <b>Pathway:</b>             | GPCR/G Protein; Metabolic Enzyme/Protease   |
| <b>Storage:</b>             | Please store the product under the recommended conditions in the Certificate of Analysis.   |

### BIOLOGICAL ACTIVITY

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|--------------------|---|
| <b>Description</b> | NN1177 is a long-acting GLP-1/glucagon receptor co-agonist. NN1177 can induce a dose-dependent body weight loss in diet-induced obese (DIO) mice <sup>[1][2]</sup> .  |
| <b>In Vitro</b>    | NN1177 (100 nM, 3 days) reduces CYP3A4 mRNA expression (57.2-71.7%) and activity (18.5-51.5%) in freshly isolated human hepatocytes <sup>[3]</sup> .<br>MCE has not independently confirmed the accuracy of these methods. They are for reference only.   |
| <b>In Vivo</b>     | NN1177 (3 or 5 nmol/kg, s.c.) induces body weight loss, loss of fat mass, and improvement in glucose tolerance in diet-induced obese (DIO) mice <sup>[1]</sup> .<br>NN1177 (0.75-4 nmol/kg, s.c., once daily, 8 weeks) reduces liver fat and inflammatory and fibrosis relevant biomarkers in C57Bl/6 mice fed a fructose and high fat rich diet (NASH model) <sup>[2]</sup> .<br>MCE has not independently confirmed the accuracy of these methods. They are for reference only. |

### REFERENCES

- [1]. Simonsen L, et al. Preclinical evaluation of a protracted GLP-1/glucagon receptor co-agonist: Translational difficulties and pitfalls. *PLoS One*. 2022 Mar 4;17(3):e0264974.
- [2]. Monfeuga T, et al. Evaluation of long acting GLP1R/GCGR agonist in a DIO and biopsy-confirmed mouse model of NASH suggest a beneficial role of GLP-1/glucagon agonism in NASH patients. *Mol Metab*. 2023 Dec 7;79:101850.
- [3]. Säll C, et al. In vitro CYP450 enzyme down-regulation by GLP-1/glucagon co-agonist does not translate to observed drug-drug interactions in the clinic. *Drug Metab Dispos*. 2022 Jun 9;DMD-AR-2022-000865.

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

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