

The 3-week NASH mouse model

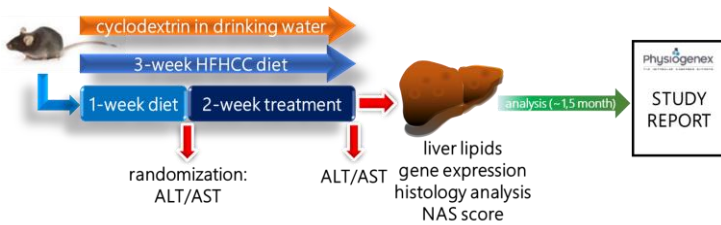
✓ A fast, costless nutritional mouse model, to rapidly evaluate your compounds targeting (NASH)

Key benefits

- ✓ **Get a deep evaluation** (biochemistry, histology and NAS score) of your compounds targeting NASH **within 2 months**
- ✓ **1 week diet-induction and 2 weeks treatment** to evaluate the impact of your drug vs. benchmarks: Our unique nutritional model develops NASH features like high ALT/AST levels, increased liver lipids, severe inflammation, and portal fibrosis **within 3 weeks**

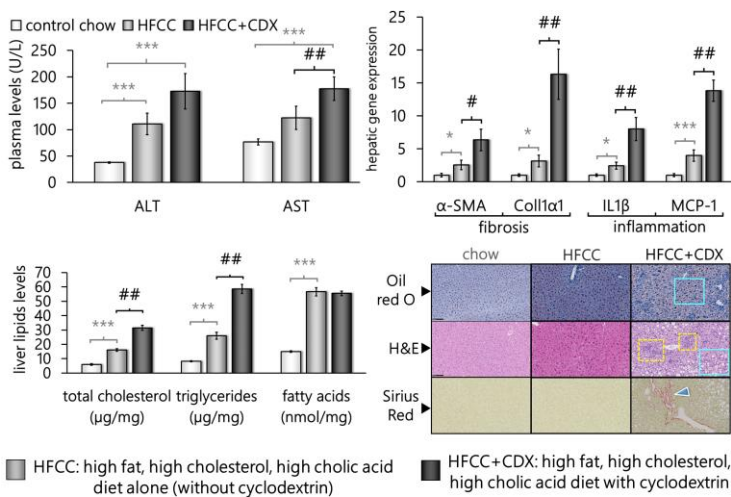
ANIMAL MODEL

- **Background strain:** C57BL/6J mouse
- **Our original diet-induced NASH:** 60% high fat diet supplemented with cholesterol/cholic acid (HFCC)+ cyclodextrin in drinking water (HFCC+CDX)
- **Study duration:** 3 weeks
- **Reference compounds:** elafibranor (dual PPAR α / δ agonist) and liraglutide (GLP-1 receptor agonist)



CHARACTERISTICS AFTER 3 WEEKS OF DIET

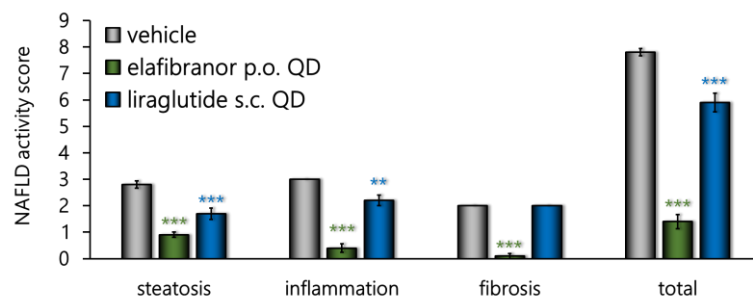
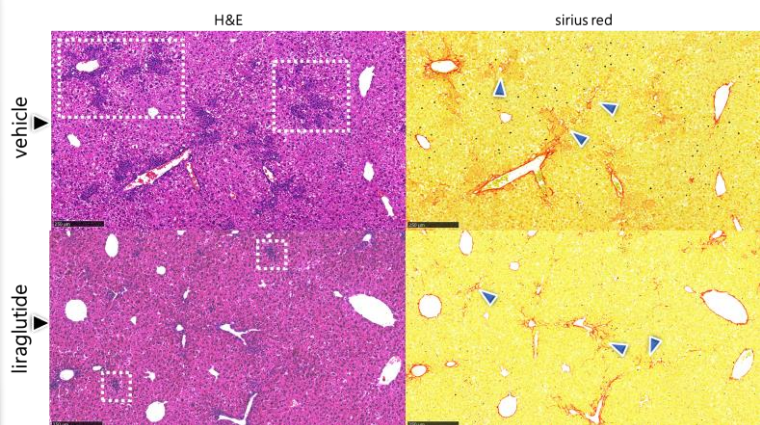
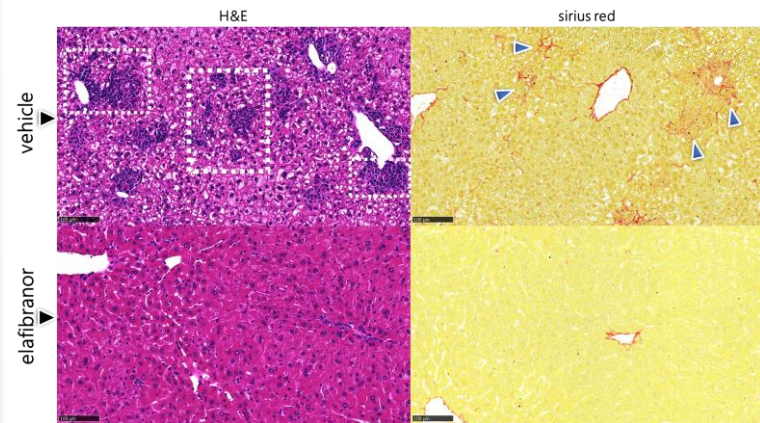
3-WEEK NASH DIET (HFCC+CDX) RESULTS LIVER LIPIDS ACCUMULATION, INFLAMMATION AND FIBROSIS



Plasma transaminases (upper left panel), hepatic gene expression (upper right panel), hepatic lipids levels (lower left panel), representative oil red O, H&E and Sirius Red staining (lower right panel) in mice fed a chow, HFCC or HFCC+CDX for 3 weeks. Blue square indicates liver steatosis, yellow dashed square inflammatory foci, blue arrows indicates fibrosis. * $p < 0.05$ and *** $p < 0.001$ vs. control chow, # $p < 0.05$, ## $p < 0.01$ and ### $p < 0.001$ HFCC+CDX vs. HFCC.

EFFECTS OF POSITIVE CONTROLS

2-WEEK TREATMENT WITH ELAFIBRANOR OR LIRAGLUTIDE IMPROVES NASH



Representative H&E and Sirius Red staining in HFCC+CDX fed mice treated for 2 weeks with vehicle or elafibranor (upper panel), vehicle or liraglutide (middle panel). White dashed squares indicate hepatic steatosis and inflammatory foci, blue arrows indicates fibrosis.

Lower panel: NAFLD activity score in HFCC+CDX fed mice treated for 2 weeks with vehicle, elafibranor or liraglutide ** $p < 0.01$ and *** $p < 0.001$ vs. vehicle.