

Effects of THR- β agonist resmetirom in the 3-week NASH mouse model

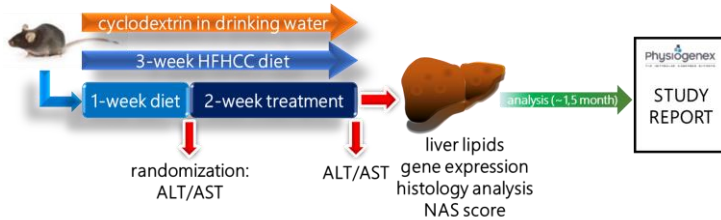
✓ A fast, costless nutritional mouse model, to rapidly evaluate your compounds targeting (NASH)

Key benefits

- ✓ **Get a complete evaluation** (biochemistry, histology and NAScore) of your compounds targeting NASH **within 2 months**. Our unique nutritional model **develops NASH features within 3 weeks**.
- ✓ **Evaluate** the impact of your drug, alone or in combination with a clinical benchmark, versus resmetirom, a THR- β agonist improving NASH and fibrosis in phase III clinical trials.

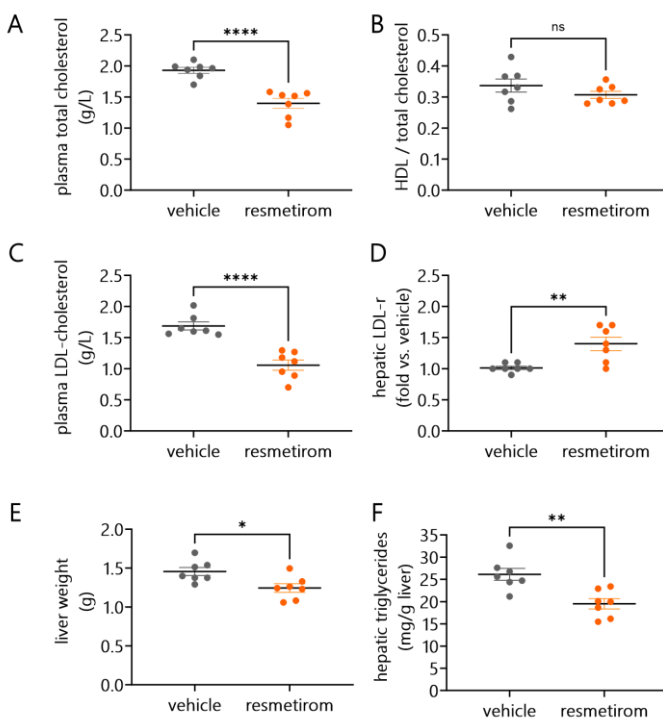
ANIMAL MODEL

- **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:** resmetirom, obeticholic acid, semaglutide, elafibranor, lanifibranor, firsocostat, etc.



EFFECTS OF RESMETIROM

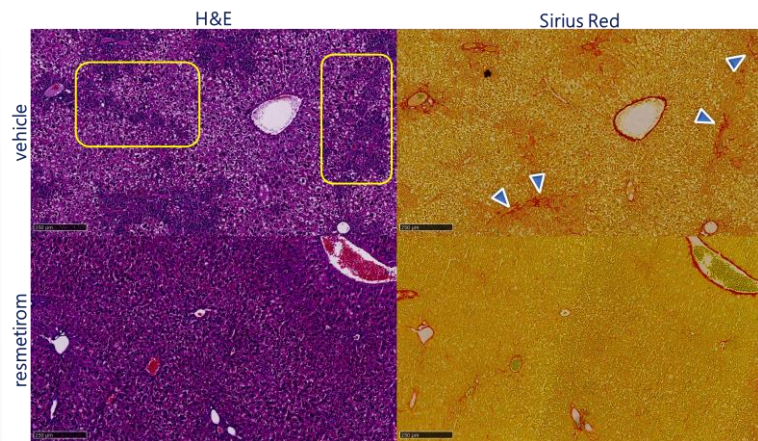
2-WEEK TREATMENT WITH RESMETIROM REDUCES LDL-C, LIVER WEIGHT AND FAT CONTENT



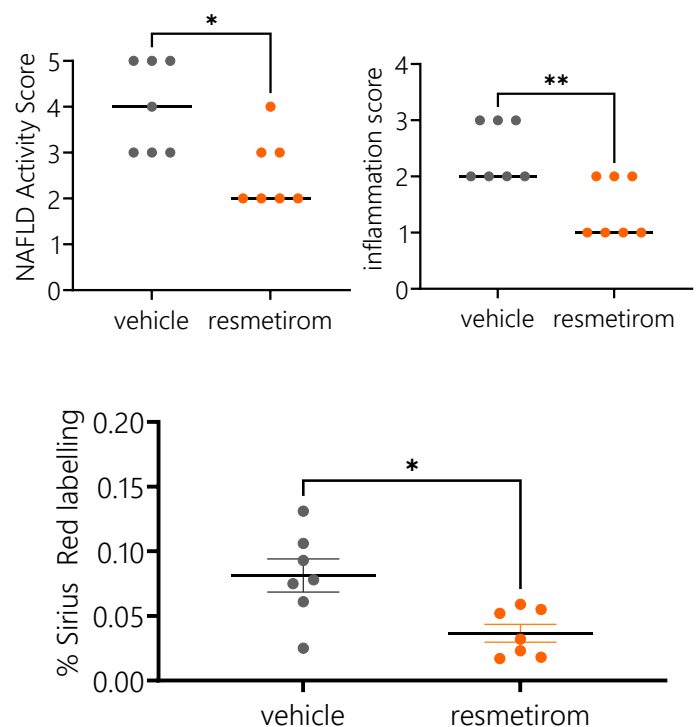
Plasma total cholesterol (A), HDL / total cholesterol ratio (B), LDL-cholesterol levels (C), hepatic LDL-r gene expression (D), liver weight (E) and hepatic triglycerides levels (F) in HFCC+CDX fed mice treated for 2 weeks with vehicle or resmetirom.

* $p < 0.05$, ** $p < 0.01$ and **** $p < 0.0001$ vs. vehicle.

2-WEEK TREATMENT WITH RESMETIROM REDUCES NAFLD ACTIVITY AND INFLAMMATION SCORES AND FIBROSIS



Representative H&E (left panel) and Sirius Red (right panel) staining in HFCC+CDX fed mice treated for 2 weeks with vehicle or resmetirom. Yellow circles indicate liver microvesicular steatosis and inflammatory foci, blue arrows indicates fibrosis.



NAFLD activity score and inflammation score (upper panel), and % Sirius Red labelling (lower panel) in HFCC+CDX fed mice treated for 2 weeks with vehicle or resmetirom.

* $p < 0.05$ vs. vehicle.