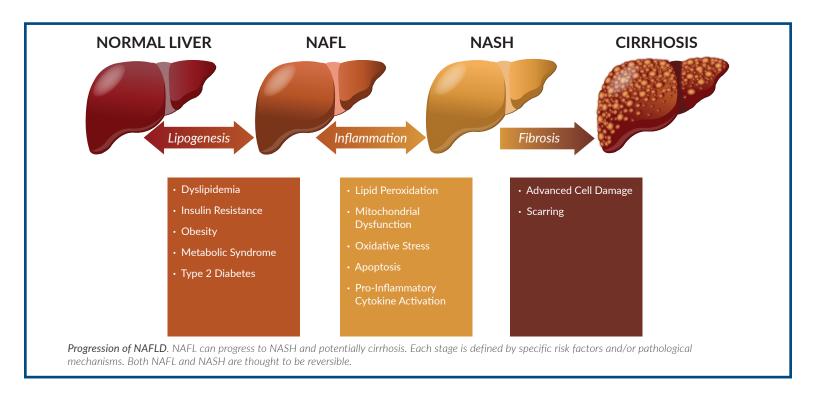
Research Tools for Fatty Liver Diseases



Non-alcoholic fatty liver disease (NAFLD) refers to a group of conditions characterized by steatosis (NAFL), the abnormal accumulation of lipids in hepatocytes that is not a consequence of excess alcohol consumption. NAFL can progress to non-alcoholic steatohepatitis (NASH), a severe form of NAFLD characterized by the presence of sustained liver inflammation and cellular damage. NASH can progress further to more chronic conditions such as cirrhosis or hepatocellular carcinoma. The molecular mechanisms involved in the development and progression of fatty liver diseases remain poorly understood. Cayman offers a range of biochemical tools and assay kits to assist investigators in their study of this growing health problem.



Products to Study Lipogenesis

NAFLD is marked by excessive fat accumulation in the liver (steatosis), which can result from systemic and hepatic desensitization of insulin signaling pathways.

Biochemicals

| Item No. | Product Name | Description |
|----------|--|---|
| 22478 | Aramchol | Prevents the formation of cholesterol crystals and gallstones in mice fed a lithogenic, high-fat diet |
| 11686 | Betulinic Acid | TGR5 agonist (EC $_{50}$ = 1 μM); reduces NF-kB signaling in LPS-treated macrophages and increases ABCA1 expression |
| 12032 | BI6015 | $\mbox{HNF4}\alpha \mbox{ antagonist that causes dose-dependent fat accumulation (steatosis) in mouse liver}$ |
| 10008335 | all-cis-4,7,10,13,16-Docosapentaenoic Acid | A DPA isomer whose concentrations increased in NASH |
| 16624 | GPR120 Compound A | Decreases hepatic steatosis when included at 30 mg/kg body weight in a high-fat diet fed to obese mice |

Biochemicals Continued

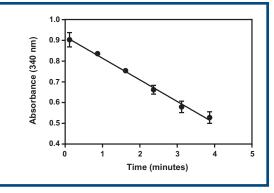
| Item No. | Product Name | Description |
|----------|---------------------------------|---|
| 17678 | INT-777 | TGR5 agonist (EC $_{50}$ = 0.82 μ M); reduces hepatic steatosis and adiposity in mice subjected to diet-induced obesity |
| 22287 | Ipragliflozin | A potent inhibitor of sodium-glucose cotransporter 2 (SGLT2; $IC_{50} = 7.4$ nM for human SGLT2 in vitro) |
| 13118 | Metformin (hydrochloride) | Improves fatty liver disease by reversing hepatic steatosis in ob/ob mice |
| 18293 | MHY908 | Improves hepatic steatosis by enhancing CPT1 levels and reducing ER stress and JNK activation in livers of db/db mice |
| 15192 | ML-261 | Inhibits hepatic lipid droplet formation (IC_{50} = 69.7 nM in mouse AML-12 hepatocytes) |
| 10007853 | Muraglitazar | A dual agonist of PPAR α and PPAR γ that improves diabetes and other metabolic abnormalities while preserving β -cell function in db/db mice |
| 14173 | Naringenin | Corrects VLDL overproduction, ameliorates hepatic steatosis, and attenuates dyslipidemia in LDL receptor-deficient mice fed a Western diet |
| 17118 | Niclosamide (ethanolamine salt) | Prevents high-fat diet-induced hepatic steatosis in db/db mice |
| 16209 | p-nitro-Pifithrin-α | Attenuates steatosis and liver injury in mice fed a high-fat diet |
| 71745 | Pioglitazone | A selective PPARγ1 agonist with antidiabetic actions (one-tenth as potent as rosiglitazone) |
| 71740 | Rosiglitazone | A potent antidiabetic agent that acts as a selective PPAR γ agonist ($K_d = 43 \text{ nM}$) |
| 18771 | SR9238 | LXR α and LXR β inverse agonist (IC $_{50}$ s = 214 and 43 nM, respectively); suppresses hepatic lipogenesis, inflammation, lipid accumulation, and fibrosis in mouse models of NASH |
| 16291 | TGR5 Receptor Agonist | Activator of TGR5 (pEC ₅₀ = 6.8-7.5); increases GLP-1 secretion and improves glucose disposal |

Convenient Monitor of ALT Levels

Alanine Transaminase Colorimetric Activity Assay Kit

Item No. 700260

- Measure ALT activity in serum, plasma, tissue samples, and cell lysates
- Assay 47 samples in duplicate
- Measure ALT activity down to 0.006 U/ml
- Plate-based colorimetric measurement (340 nm)



Additional Assay Kits

| Item No. | Product Name | Description |
|----------|---|---|
| 10007640 | Cholesterol Fluorometric Assay Kit | Quantitation of total cholesterol in plasma or serum |
| 700310 | Free Fatty Acid Fluorometric Assay Kit | Measure FFAs in plasma, serum, and urine |
| 15741 | FXR (human) Reporter Assay Kit | Screen test samples to quantify functional activity, either agonist or antagonist, that they may exert against human FXR |
| 10011125 | LDL Uptake Cell-Based Assay Kit | Measure LDL uptake in cultured cells |
| 500010 | Leptin (human) EIA Kit (Manufactured by Bertin Pharma) | Measure leptin in human serum, plasma, and culture supernatant samples Also available: Leptin (mouse/rat) EIA Kit (Item No. 10007609) and Leptin Receptor (human) EIA Kit (Item No. 10007608) |
| 500001 | Lipid Droplets Fluorescence Assay Kit | Detect lipid droplets, cellular organelles that are also referred to as lipid bodies, oil bodies, or adiposomes |
| 10010854 | SREBP-1 Transcription Factor Assay Kit | A sensitive, non-radioactive method of detecting SREBP-1 from whole cell lysates |
| 10012643 | Steatosis Colorimetric Assay Kit | Detect excessive lipid accumulation in cells |
| 601440 | TGR5 (GP-BAR1) Reporter Assay Kit | Screen for TGR5 agonists, antagonists, and modulators |
| 10010303 | Triglyceride Colorimetric Assay Kit | Measure TG levels in plasma, serum, cell lysates, and tissue homogenates |

Products to Study Inflammation

NASH is marked by inflammation of the liver (steatohepatitis), which can result from cytokine dysregulation, oxidative stress, and hepatocyte apoptosis.

Biochemicals

| Item No. | Product Name | Description |
|----------|------------------------------|---|
| 70935 | (–)-Epigallocatechin Gallate | Inhibits cellular oxidation of LDLs |
| 14127 | Febuxostat | Inhibits oxidized and reduced forms of xanthine oxidase |
| 10739 | Isoliquiritigenin | Exhibits antioxidant, anti-inflammatory, and antitumor activities, as well as hepatoprotection against steatosis-induced oxidative stress |
| 11948 | MN-001 | Produces antifibrotic and anti-inflammatory activity in models of NASH |
| 16093 | Myrtillin | A natural anthocyanin with antioxidant activity |
| 32100 | 4-hydroxy Nonenal | A marker of lipid peroxidation |
| 18720 | Pentoxifylline | Anti-inflammatory activity, inhibiting TNF-α production |
| 70675 | trans-Resveratrol | Phenolic antioxidant with anti-inflammatory activity |
| 18771 | SR9238 | LXR α and LXR β inverse agonist (IC $_{50}$ s = 214 and 43 nM, respectively); suppresses hepatic lipogenesis, inflammation, lipid accumulation, and fibrosis in mouse models of NASH |
| 10011659 | Trolox | Derivative of vitamin E with potent antioxidant properties |

Convenient Monitor of the TNF-a Response

TNF-α (human) ELISA Kit

Item No. 589201

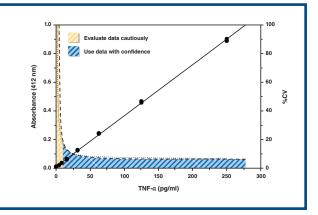
• Measure TNF- α from human plasma, serum, and other sample matrices

• Assay Range: 3.9-250 pg/ml with an LOD of 3.9 pg/ml

• Incubation: 18 hours | Development: 60-90 minutes

• Read: Colorimetric at 405-420 nm

Also available: TNF- α (mouse) ELISA Kit (Item No. 500850)



Additional Assay Kits

| Item No. | Product Name | Description |
|----------|---|---|
| 701080 | COX-2 (human) Inhibitor Screening Assay Kit | Complete Drug Discovery Assay for screening COX-2 inhibitors |
| 702000 | COX-2 (ovine) Inhibitor Screening Assay Kit | Complete Drug Discovery Assay for screening COX-2 inhibitors |
| 10011236 | C-Reactive Protein (human) ELISA Kit | Measure CRP from human plasma without prior sample purification |
| 501030 | Interleukin-6 (human) ELISA Kit | Measure IL-6 from human plasma, serum, synovial fluid, and other sample matrices |
| 600610 | Neutrophil Elastase Activity Assay Kit | Measure neutrophil elastase released by neutrophils after various experimental conditions |
| 601290 | ROS Detection Cell-Based Assay Kit (DHE) | Measure superoxide and hydrogen peroxide levels in living cells |
| 10009055 | TBARS Assay Kit | Measure lipid peroxidation in plasma, serum, urine, tissue homogenates, and cell lysates |
| 700870 | TBARS (TCA Method) Assay Kit | Measure lipid peroxidation in plasma, serum, urine, tissue homogenates, and cell lysates; includes TCA precipitation procedure for improved sample processing |

Additional lipid peroxidation, mitochondrial dysfunction, oxidative stress, and apoptosis assay kits available online

Products to Study Fibrosis

Cirrhosis is marked by prolonged cell damage and fibrosis, which results from the activation of hepatic stellate cells and the emergence of fibrotic progenitors.

Biochemicals

| Item No. | Product Name | Description |
|----------|-----------------------------|---|
| 16018 | Camostat (mesylate) | A protease inhibitor that disrupts the proliferation of stellate cells |
| 10008199 | Linolenic Acid ethyl ester | Stimulates intracellular signaling pathways in hepatic stellate cells |
| 11948 | MN-001 | Produces antifibrotic and anti-inflammatory activity in models of NASH |
| 17278 | Necrox-5 (methanesulfonate) | A necrosis inhibitor that protects cells from carbon tetrachloride-induced liver injury and chronic liver fibrosis in rodent models |
| 19131 | NK 252 | Exhibits antifibrotic effects in a rat model of NASH |
| 13986 | Pirfenidone | Antioxidant, anti-inflammatory, and antifibrotic effects |
| 18771 | SR9238 | LXR α and LXR β inverse agonist (IC $_{50}$ S = 214 and 43 nM, respectively); suppresses hepatic lipogenesis, inflammation, lipid accumulation, and fibrosis in mouse models of NASH |

Proteins

| Item No. | Product Name | Description |
|----------|---------------------------|----------------------------------|
| 10009547 | FABP1 (human recombinant) | Potential marker of liver injury |
| 10005200 | FABP1 (rat recombinant) | Potential marker of liver injury |

Nuclear Receptor Assays



Current NASH research indicates drug and treatment discovery relies on nuclear receptor activation, specifically the CAR, FXR, LXR α , LXR β , PPAR α , PPAR γ , PPAR β / δ , PXR, ROR γ , and VDR receptors. Cayman partners with Indigo Biosciences to offer assay kits for these NAFLD/NASH-associated nuclear receptors.

| Item No. | Product Name |
|----------|----------------------------------|
| 15748 | CAR1 (human) Reporter Assay Kit |
| 15747 | CAR3 (human) Reporter Assay Kit |
| 15741 | FXR (human) Reporter Assay Kit |
| 20349 | FXR (mouse) Reporter Assay Kit |
| 15735 | LXRα (human) Reporter Assay Kit |
| 15734 | LXRβ (human) Reporter Assay Kit |
| 15730 | PPARα (human) Reporter Assay Kit |
| 15762 | PPARα (mouse) Reporter Assay Kit |
| 15764 | PPARα (rat) Reporter Assay Ki |
| 15729 | PPARγ (human) Reporter Assay Kit |

| 15761 PPARγ (mouse/rat) Reporter Assay Kit 15731 PPARδ (human) Reporter Assay Kit 15763 PPARδ (mouse) Reporter Assay Kit 15765 PPARδ (rat) Reporter Assay Kit 15732 PPAR (human) Reporter Assays Panel 17528 Pregnane X (human) Receptor Assay Kit 17585 Pregnane X (rat) Receptor Assay Kit | Item No. | Product Name |
|--|----------|---------------------------------------|
| 15763 PPARδ (mouse) Reporter Assay Kit 15765 PPARδ (rat) Reporter Assay Kit 15732 PPAR (human) Reporter Assays Panel 17528 Pregnane X (human) Receptor Assay Kit | 15761 | PPARγ (mouse/rat) Reporter Assay Kit |
| 15765 PPARδ (rat) Reporter Assay Kit 15732 PPAR (human) Reporter Assays Panel 17528 Pregnane X (human) Receptor Assay Kit | 15731 | PPARδ (human) Reporter Assay Kit |
| 15732 PPAR (human) Reporter Assays Panel 17528 Pregnane X (human) Receptor Assay Kit | 15763 | PPARδ (mouse) Reporter Assay Kit |
| 17528 Pregnane X (human) Receptor Assay Kit | 15765 | PPARδ (rat) Reporter Assay Kit |
| | 15732 | PPAR (human) Reporter Assays Panel |
| 17585 Pregnane X (rat) Recentor Assay Kit | 17528 | Pregnane X (human) Receptor Assay Kit |
| 1, 555 1.166, nation (i.e., necespter, neces) the | 17585 | Pregnane X (rat) Receptor Assay Kit |
| 15757 RORγ (human) Reporter Assay Kit | 15757 | RORγ (human) Reporter Assay Kit |
| 15742 VDR (human) Reporter Assay Kit | 15742 | VDR (human) Reporter Assay Kit |

To view a complete list of our NAFLD research tools, visit us online at www.caymanchem.com