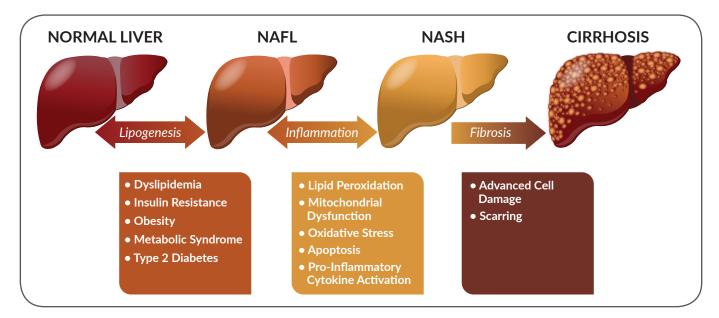
Research Tools for Fatty Liver Diseases

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 nonalcoholic 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.







Progression of NAFLD. NAFL can progress to NASH and potentially cirrhosis. Each stage is defined by specific risk factors and/or pathological mechanisms. Both NAFL and NASH are thought to be reversible.

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- κB signaling in LPS-treated macrophages and increases ABCA1 expression
12032	BI6015	$HNF4\alpha \text{ antagonist that causes dose-dependent fat accumulation (steatosis) in mouse liver}$
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

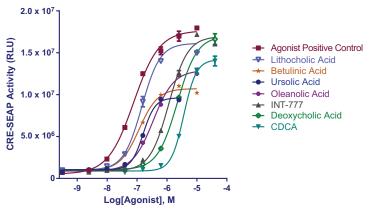
Item No.	Product Name	Description
22287	Ipragliflozin	A potent inhibitor of sodium-glucose cotransporter 2 (SGLT2; $IC_{50} = 7.4$ nM for human SGLT2 in vitro)
13118	Metformin (hydrochloride)	Improves fatty liver disease by reversing hepatic steatosis in ob/ob mice
18293	MHY908	Improves hepatic steatosis 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 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

Screen for Modulators of a Bile Acid Receptor

TGR5 (GP-BAR1) Reporter Assay

Item No. 601440

- Reverse transfection reporter assay to screen for TGR5 agonists, antagonists, and modulators
- No stable cell line required, simply add regular HEK293 or HEK293T cells to the plate the day before assaying
- 96-Well strip format for flexible plate map or assay on a partial plate
- Positive control agonist included
- Obtain results within 24-30 hours after adding cells



This assay has been thoroughly validated with seven known TGR5 receptor agonists in addition to the TGR5 Receptor Agonist Positive Control.

Known TGR5 Agonists

Item No.	Product Name
11686	Betulinic Acid
10011286	Chenodeoxycholic Acid
20756	Deoxycholic Acid
17678	INT-777
20253	Lithocholic Acid
11726	Oleanolic Acid
16291	TGR5 Receptor Agonist
10072	Ursolic Acid

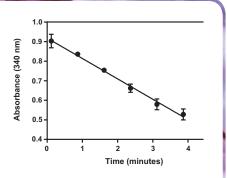
Additional bile acids and derivatives available online

Additional Bile Acids

Item No.	Product Name
20250	Cholic Acid
20274	Glycodeoxycholic Acid
21723	Glycolithocholic Acid
20294	Hyodeoxycholic Acid
20275	Taurchenodeoxycholic Acid (sodium salt)
15935	Taurodeoxycholic Acid (sodium salt)
20277	Tauroursodeoxycholic Acid
15121	Ursideoxycholic Acid (sodium salt)

Convenient Assay to Monitor 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

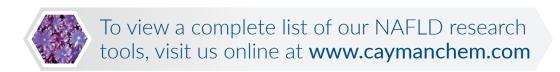
Item No.	Product Name	Description
10007640	Cholesterol Fluorometric Assay Kit	Quantitation of total cholesterol in plasma or serum
700310	Free Fatty Acid Fluorometric Assay Kit	Measure FFAs in plasma, serum, and urine
15741	FXR (human) Reporter Assay Kit	Screen test samples to quantify functional activity, either agonist or antagonist, that they may exert against human FXR
10011125	LDL Uptake Cell-Based Assay Kit	Measure LDL uptake in cultured cells
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
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-α Response

TNF-α (human) ELISA Kit

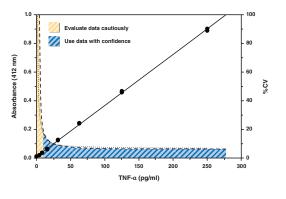
Item No. 589201

• 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

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Tel: +33 (0)139 306 036 Email: tech@bertin-bioreagent.com Web: www.bertin-bioreagent.com