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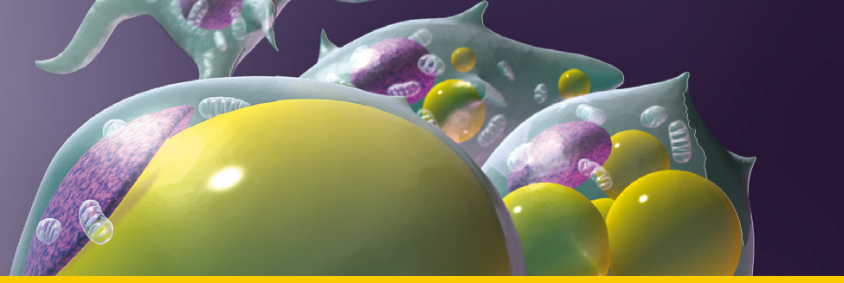
LIFE SCIENCES



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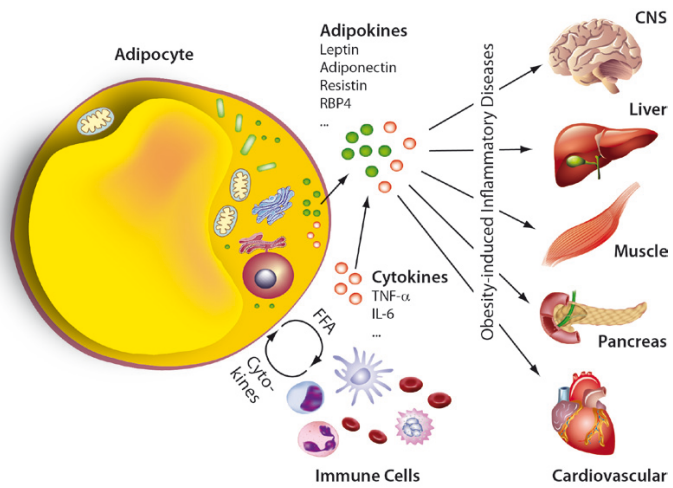
3rd Edition

Obesity & Diabetes Research

Focus: White & Brown Fat Cells as Endocrine Tissues

Two major types of adipose tissue exist in mammals, named white (WAT) and brown adipose tissue (BAT) composed mainly of white (see below) or brown adipocytes (see page 8), respectively. **White adipose tissue (WAT)** is found throughout the body, primarily under the skin (subcutaneous fat) as well as in larger deposits in the abdomen (visceral fat). White adipocytes act as storage cells for neutral triacylglycerols, storing excess calories for use in times of scarcity. WAT contributes to whole body insulation and actively communicates with key organs to maintain metabolic homeostasis by secreting adipokines.

Adipokines are defined generally as biologically active substances produced in white adipose tissue (WAT) that act in an autocrine/paracrine or endocrine fashion and communicate with the brain, heart, vasculature, liver and muscle. Some adipokines are produced exclusively or predominantly by adipose tissue, whereas others may be produced in a variety of different tissues. The diversity of the adipokines is considerable, in terms of both, protein structure and function. Adipokines include classical cytokines (e.g. TNF-α, IL-6), chemokines (e.g. MCP-1), proteins of the alternative complement system (e.g. Adipsin), proteins involved in vascular hemostasis (e.g. PAI-1), the regulation of blood pressure (Angiotensinogen), lipid metabolism (e.g. RBP4), glucose homeostasis (e.g. Adiponectin, Leptin, Progranulin, Nampt/Visfatin/PBEF, Resistin, Vaspin, Omentin, Lipocalin-2, Apelin, DPP-4, CTRPs, selected ANGPTLs), angiogenesis (e.g. VEGF, NGF) and lipid mobilization (Zinc-α-2-glycoprotein). Adipokines have either pro-inflammatory or anti-inflammatory activities and exhibit a wide range of functions including the regulation of food intake and body weight homeostasis, insulin sensitivity, cell proliferation and angiogenesis, immunity, inflammation or vascular homeostasis. During obesity (see page 4), adipokines are dysregulated and create a state of **chronic low-grade inflammation** responsible for the different obesity-linked pathologies and the onset of insulin resistance. Although **brown adipose tissue (BAT)** also produces adipokines (see page 8), the endocrine role of BAT in metabolic diseases is not fully investigated. A growing interest in adipokines and myokines as biomarkers of low-grade inflammation and metabolic diseases emerges.



SELECTED REVIEW ARTICLE

Adipose tissue: between the extremes: A. Vegiopoulos, et al.; EMBO J. **36**, 1999 (2017)

FIGURE: Schematic interaction between adipocytes and immune cells.

*Adapted from H. Cao; J. Endocrinol. **220**, T47 (2014)*

CONTENTS

KEY Adipokines: Novel Biomarkers and Regulators of Diabetes, Obesity, Insulin Resistance and Inflammation

Adiponectin

Adiponectin is an important adipocyte-derived anti-inflammatory hormone that regulates metabolism of lipids and glucose. Its receptors (AdipoR1, AdipoR2, T-cadherin) appear to exert actions in peripheral tissues by activating the AMP-activated protein kinase, p38-MAPK, PPARα and NF-κB and exerting a wide range of beneficial physiological actions, including antidiabetic, anti-inflammatory, anti-atherosclerotic and cardioprotective effects. Adiponectin is the most abundant adipokine in the circulation and its levels are substantially altered in obesity, type 2 diabetes, cardiovascular disease, nonalcoholic fatty liver disease (NAFLD), obesity-related inflammation and various cancers.

ELISA KITS / THE STANDARDS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Adiponectin (human) ELISA Kit	AG-45A-0001Y	96 wells | 2 x 96 wells	100 pg/ml	0.5 to 32 ng/ml	C, P, S, U
Adiponectin (mouse) ELISA Kit	AG-45A-0004Y	96 wells | 2 x 96 wells	50 pg/ml	0.125 to 8 ng/ml	C, P, S
Adiponectin (rat) ELISA Kit	AG-45A-0005Y	96 wells | 2 x 96 wells	50 pg/ml	0.375 to 24 ng/ml	C, P, S
RECOMBINANT PROTEINS	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
Adiponectin (human) (rec.)	AG-40B-0030	50 µg | 500 µg	HEK 293 cells	<0.01EU/µg	Hu
Adiponectin (mouse) (rec.)	AG-40B-0026	50 µg | 500 µg	HEK 293 cells	<0.01EU/µg	Ms

Nampt [Visfatin; PBEF]

Nicotinamide phosphoribosyltransferase (NAMPT) is a regulator of the intracellular NAD pool. Through its NAD-biosynthetic activity, NAMPT influences the activity of NAD-dependent enzymes, thereby regulating cellular metabolism. In addition to its enzymatic function, extracellular NAMPT (also called Visfatin or PBEF1) has cytokine-like activity. Altered levels are associated with various metabolic disorders, including obesity, nonalcoholic fatty liver disease (NAFLD) and type 2 diabetes by influencing the oxidative stress response, apoptosis, lipid and glucose metabolism, inflammation and insulin resistance. NAMPT plays a crucial role in cancer cell metabolism and is often overexpressed in tumor tissues, making it an attractive therapeutic cancer drug target.

ELISA KITS / THE STANDARDS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Nampt (human) ELISA Kit	AG-45A-0006Y	96 wells | 2 x 96 wells	30 pg/ml	0.125 to 8 ng/ml	S
Nampt (human) (IntraCellular) ELISA Kit	AG-45A-0008Y	96 wells | 2 x 96 wells	30 pg/ml	0.25 to 16 ng/ml	L
Nampt (mouse/rat) Dual ELISA Kit	AG-45A-0007Y	96 wells | 2 x 96 wells	50 pg/ml	0.5 to 32 ng/ml	S
RECOMBINANT PROTEINS	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
Nampt (human) (rec.)	AG-40A-0031Y	10 µg | 3 x 10 µg	HEK 293 cells	<0.01EU/µg	Hu
Nampt (mouse) (rec.)	AG-40A-0056Y	10 µg | 3 x 10 µg	CHO cells	<0.01EU/µg	Ms
POTENT INHIBITORS	PID	SIZE	***From The Manufacturer BULK AVAILABLE***		
CHS-828	AG-CR1-0064	5 mg | 25 mg			
FK-866	AG-CR1-0011	1 mg | 5 mg			

Retinol-binding Protein 4 [RBP4]

The physiological role of RBP4 is transport of retinol from the liver to peripheral tissues. RBP4 is produced in hepatocytes and adipocytes. Excessive visceral fat accumulation, followed by the development of inflammation and consequently a hormonal adipose tissue dysfunction is in direct relation with excessive RBP4 expression, orchestrated by GLUT4. Circulating RBP4 inhibits the signal pathways stimulated by insulin in skeletal muscle cells, resulting in the development of insulin resistance. Altered levels are associated with various metabolic disorders, including obesity, cardiovascular disease, nonalcoholic fatty liver disease (NAFLD) and type 2 diabetes.

ELISA KITS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
RBP4 (human) ELISA Kit (Quantitative)	AG-45A-0035Y	96 wells | 2 x 96 wells	380 pg/ml	0.39 to 25 ng/ml	C, P, S, U
RBP4 (human) Competitive ELISA Kit	AG-45A-0010Y	96 wells | 2 x 96 wells	1 ng/ml	0.001 to 5 µg/ml	C, P, S, U
RBP4 (mouse/rat) Dual ELISA Kit	AG-45A-0012Y	96 wells | 2 x 96 wells	60 pg/ml	0.188 to 12 ng/ml	C, S, U

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SAMPLES: C: Cell Culture Supernatant, **P:** Plasma, **S:** Serum; **U:** Urine; **L:** Cell Lysate; **F:** CSF
APPLICATIONS: FUNC: Functional Application; **ICC:** Immunocytochemistry; **IHC:** Immunohistochemistry; **IP:** Immunoprecipitation; **WB:** Western blot
FORMULATION: PF = Preservative free
SPECIES: Dg = Dog; **Hu** = Human; **Mo** = Monkey; **Ms** = Mouse; **Rt** = Rat; **Rb** = Rabbit

Progranulin [PGRN]

Progranulin (PGRN) is a cysteine rich secreted protein, expressed in epithelial cells, immune cells, neurons and adipocytes. PGRN was first identified as a growth factor and recently characterized as an adipokine implicated in obesity, insulin resistance and rheumatic disease. At a central level, PGRN acts as a neurotropic and neuroprotective factor and protects from neural degeneration. PGRN has pleiotropic actions and participates in several processes, such as inflammation or tumorigenesis.

Tag-free Progranulins

- **Higher activity compared to tagged Progranulins**
- **Suitable for *in vitro* and *in vivo* studies**
- **Reflects the native sequence with no additional amino acids**
- **Affinity purified**
- **Low endotoxin levels (<0.01 EU/µg)**

ELISA KITS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Progranulin (human) ELISA Kit	AG-45A-0018Y	96 wells | 2 x 96 wells	32 pg/ml	0.063 to 4 ng/ml	C, P, S, U
Progranulin (mouse) ELISA Kit	AG-45A-0019Y	96 wells | 2 x 96 wells	60 pg/ml	0.125 to 8 ng/ml	C, S
Progranulin (rat) ELISA Kit	AG-45A-0043Y	96 wells | 2 x 96 wells	40 pg/ml	0.063 to 4 ng/ml	C, S
RECOMBINANT PROTEINS	**PID**	**SIZE**	**SOURCE**	**ENDOTOXIN**	**SPECIES**
Progranulin (human) (rec.) (untagged)	AG-40A-0188Y	10 µg | 50 µg	HEK 293 cells	<0.01EU/µg	Hu
Progranulin (mouse) (rec.) (untagged)	AG-40A-0189Y	10 µg | 50 µg	HEK 293 cells	<0.01EU/µg	Ms
Progranulin (rat) (rec.) (untagged)	AG-40A-0196Y	10 µg | 50 µg	HEK 293 cells	<0.01EU/µg	Rt

Vaspin [Visceral Adipose Tissue-derived Serpin; Serpin A12]

Vaspin, a serine protease inhibitor, is an insulin-sensitizing adipokine that has been isolated from both visceral and subcutaneous white adipose tissue. Vaspin is suggested to regulate immune responses and inflammation and was found to be correlated with various metabolic parameters. Vaspin represents a novel biomarker for obesity and impaired insulin sensitivity and might serve as a new therapeutic target of metabolic syndrome diseases, such obesity-related insulin resistance and inflammation.

ELISA KITS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Vaspin (human) ELISA Kit	AG-45A-0017Y	96 wells | 2 x 96 wells	12 pg/ml	0.016 to 1 ng/ml	C, P, S
RECOMBINANT PROTEINS	**PID**	**SIZE**	**SOURCE**	**ENDOTOXIN**	**SPECIES**
Vaspin (human) (rec.)	AG-40A-0064Y	10 µg | 3 x 10 µg	HEK 293 cells	<0.01EU/µg	Hu
Vaspin (mouse) (rec.)	AG-40A-0094	10 µg | 50 µg	HEK 293 cells	<0.1EU/µg	Ms
ANTIBODIES	**PID**	**SIZE**	**ISOTYPE/SOURCE**	**APPLICATION**	**SPECIES**
anti-Vaspin (human), mAb (VP63)	AG-20A-0045	50 µg | 100 µg	Mouse IgG1κ	IHC, WB	Hu
anti-Vaspin (mouse), pAb	AG-25A-0075	100 µg	Rabbit	WB	Ms

Zinc-α-2-glycoprotein [ZAG]

Zinc-α-2-glycoprotein (ZAG) is expressed in the major white fat depots and in the interscapular brown fat of mice defining it as an adipokine. ZAG has been shown to stimulate lipolysis in *in vitro* and *in vivo* experiments. Data from genetic studies suggest that ZAG may be a candidate gene for body weight regulation. ZAG is up-regulated in urine from diabetic patients and is reported to be associated with several diseases, such as cancers, metabolic syndrome and acute sepsis.

ELISA KITS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Zinc-α-2-glycoprotein (human) TurboELISA™ Kit	AG-48B-1000	96 wells | 2 x 96 wells	0.23 ng/ml	0.9375 to 60 ng/ml	C, P, S
Zinc-α-2-glycoprotein (human) Matched Pair Detection Set	AG-46B-0008	5 x 96 wells	100 pg/ml	0.0156 to 1 ng/ml	C, P, S
RECOMBINANT PROTEIN	**PID**	**SIZE**	**SOURCE**	**ENDOTOXIN**	**SPECIES**
Zinc-α-2-glycoprotein (human) (rec.)	AG-40B-0146	10 µg | 50 µg | 3 x 50 µg	E. coli	<0.1EU/µg	Hu

A Complete Panel of Adiponectin, Nampt, RBP4, Progranulin, Vaspin and ZAG Proteins & Antibodies is available on www.adipogen.com

Obesity & Immunometabolism

During **obesity,** excess fat accumulates in adipose tissue leading to low-grade chronic inflammation. Obesity is a major risk factor for many metabolic diseases, especially diabetes and cardiovascular diseases, increasing the risk of hypertension, hyperglycemia and dyslipidemia, recognized as the **metabolic syndrome**. Obesity is also linked to a broad spectrum of pathological disorders including neurodegenerative diseases, airway disorders and cancer.

Dysregulation in **adipokines secretion**, adipocyte mitochondrial dysfunction, alteration in the gut microbiota composition are among factors involved in the development of obesity and its associated metabolic disorders. During obesity, a modulation of immune cells is observed (see below section immunometabolism and Figure). In **lean healthy adipose tissue**, Th2 cells and eosinophils secrete Th2 cytokines IL-4, IL-10 and IL-13 leading to an anti-inflammatory macrophage M2 phenotype, ensuring tissue remodelling. In **obese adipose tissue**, overnutrition leads to bigger adipocytes, which coupled with various cellular stress consequently leads to the recruitment of different immune cells and the development of a pro-inflammatory environment.

Immunometabolism describes the ability of the immune system to communicate and coordinate systemic metabolic homeostasis. Immunometabolism can be studied at macroscopic level, the whole-body metabolism and at microscopic level, the cellular bioenergetics of immune cells. Adipose tissue illustrates best the interdependency of both arms of immunometabolism (whole-body metabolism and the microscopic metabolism) and provides examples of changes in both the lean and obese states (see Figure). Lean adipose tissue is characterized by an enrichment of immune cells whose phenotype and cytokine profiles maintain a state of type 2 immunity necessary for the health of the tissue. Obesity is characterized by an accumulation of inflammatory immune cells and loss of protective lymphocytes due to change in the composition of fatty acids, glucose and oxygen availability that may provide different metabolic substrates to immune cells and adipocytes.

SELECTED REVIEWS: Adipose tissue: between the extremes: A. Vegiopoulos, et al.; EMBO J. **36,** 1999 (2017) • Adipose tissue at the nexus of systemic and cellular immunometabolism: A.C. Kohlgruber, et al.; Sem. Immunol. **28,** 431 (2016)

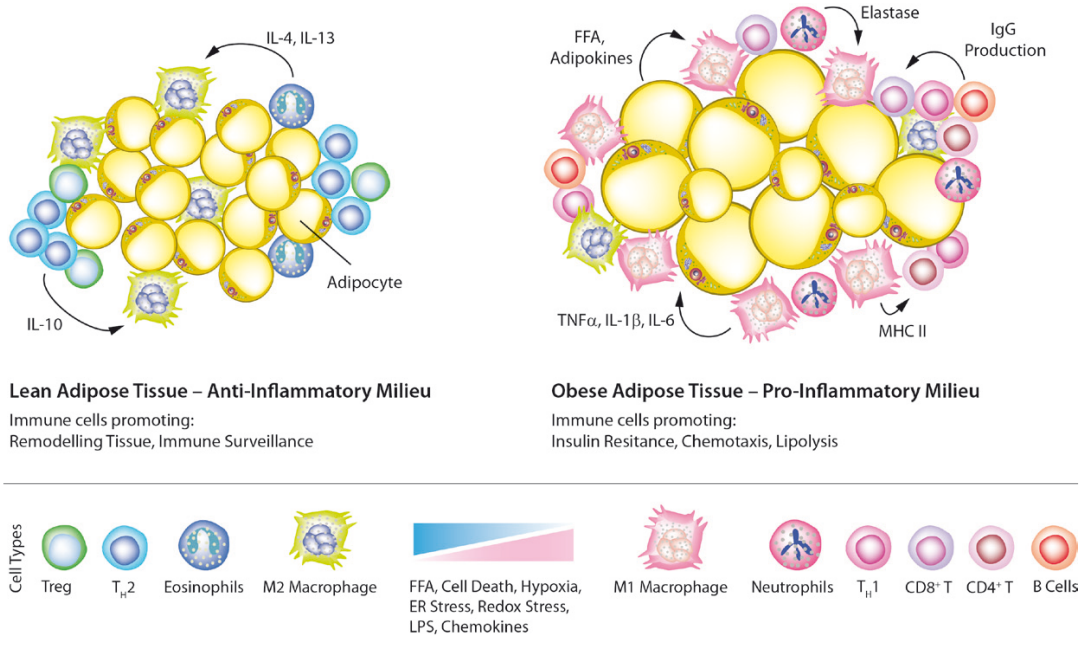


FIGURE: Modulation of immunometabolism during obesity.

*Adapted from H.L. Kammoun, et al.; Rev. Endocr. Metab. Disord. **15,** 31 (2014)*

Other Obesity-related Proteins & Antibodies

PROTEINS	PID
Calreticulin (human) (rec.) (His)	AG-40A-0132
Clusterin (secretory form) (human) (rec.)	AG-40A-0050Y
Clusterin (nuclear form) (human) (rec.) (His)	AG-40A-0047
Clusterin (nuclear form) (mouse) (rec.) (His)	AG-40A-0057
CREB-binding Protein (mouse) (rec.) (His)	AG-40T-0016
NEW CTHRC1 (human) (rec.)	AG-40B-0157
NEW CTHRC1 (mouse) (rec.)	AG-40B-0154
FABP1 (human) (rec.) (His)	AG-40A-0039T
FABP3 (human) (rec.) (untagged)	AG-40B-6002
FABP4 (human) (rec.) (His)	AG-40A-0035
FTO (human) (rec.) (His)	AG-40A-0112
FTO (mouse) (rec.) (His)	AG-40A-0127
NEW IDO (human) (rec.) (His) (highly active)	AG-40B-0161
Lipocalin-2 (human) (rec.)	AG-40B-6001
NAD Kinase (human) (rec.) (His) (highly active)	AG-40T-0091
NMNAT1 (human) (rec.) (His) (highly active)	AG-40T-0092
NMNAT3 (human) (rec.) (His) (highly active)	AG-40T-0093
Omentin (human) (rec.)	AG-40B-0042
PEDF (human) (rec.)	AG-40B-0077
PEDF (mouse) (rec.)	AG-40B-0118
Resistin (human) (rec.)	AG-40A-0010Y
Resistin (mouse) (rec.)	AG-40A-0011

ANTIBODIES	PID
Calreticulin (human), mAb (CR213-2AG)	AG-20A-0079
Calreticulin (human), pAb	AG-25A-0094
Clusterin (human), pAb	AG-25A-0099
Clusterin (mouse), pAb	AG-25A-0054
FABP3 (human), pAb	AG-25A-0040
FABP4 (human), pAb	AG-25A-0041
FTO (human), mAb (AG103)	AG-20A-0092
FTO (mouse), mAb (FT62-6)	AG-20A-0083
IDO (human), mAb (ID 177)	AG-20A-0035
IDO (mouse), pAb	AG-25A-0032
Leptin (rat), mAb (RLEP 227)	AG-20A-0018
NEW MPC-2, mAb (JCM-1)	AG-20B-0071
NMNAT2 (human), mAb (Nady-1)	AG-20A-0087
Obestatin (human), pAb	AG-25A-0043
NEW PEDF (human), mAb (rec.) (Serpy-1-4)	AG-27B-0014
RELM-α (mouse), mAb (MREL 384)	AG-20A-0020
RELM-β (mouse), mAb (MRB 46L)	AG-20A-0026
Resistin (human), pAb	AG-25A-0013
Resistin (mouse), mAb (MRES 06)	AG-20A-0004
Resistin (rat), mAb (RRES 07)	AG-20A-0015
Stearoyl-CoA Desaturase-1 (mouse), pAb	AG-25A-0031
TDO (human), pAb	AG-25A-0106
TRB-3 (human), pAb	AG-25A-0059

BULK

IBMX

Enhances Differentiation of 3T3-L1 Cells

IBMX [3-Isobutyl 1-methylxanthine]

AG-CR1-3512-M500	500 mg
AG-CR1-3512-G001	1 g

Formula: $C_{10}H_{14}N_4O_2$
MW: 222.3
CAS: 28822-58-4

BULK

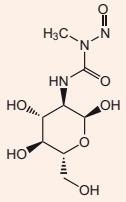
Streptozotocin

STANDARD Diabetes Inducer

Streptozotocin

AG-CN2-0046-M050	50 mg
AG-CN2-0046-M250	250 mg
AG-CN2-0046-G001	1 g

Formula: $C_8H_{15}N_3O_7$
MW: 265.2
CAS: 18883-66-4



IL-33 and Adipose Tissue Homeostasis

Lean adipose tissue contains adipocytes, regulatory immune cells and adipose stroma that contribute to fat tissue homeostasis. Adipocytes of lean tissue secrete adipokines (e.g. adiponectin, an anti-inflammatory protein), which play important roles in the regulation of systemic metabolism (immunometabolism) and have a profound impact on immune cell behavior. Various immune cells are implicated in lean adipose tissue remodeling, such as invariant-chain natural killer T (iNKT) cells, eosinophils, type 2 innate lymphocytes ILC2s and regulatory T cells (Tregs). These immune cells maintain homeostasis, preserving insulin sensitivity and glucose tolerance and keeping adipose tissue macrophages (ATMs) in an anti-inflammatory, M2-like state [1] (see Figure).

During high-fat diet and obesity, fat cells increase (hypertrophy) producing less adiponectin and more pro-inflammatory molecules such as leptin, IL-6 and monocyte chemo-attractant protein-1 (MCP-1). Inflammatory immune cells such as neutrophils or NK cells detect adipose stress, accumulate and secrete IFN-γ, driving pro-inflammatory M1 macrophage differentiation leading to a chronic inflammatory state.

IL-33, a cytokine abundantly expressed by adipose tissue stroma, is of particular importance for adipose homeostasis. Although upon infection and allergy, IL-33 is classified as a pro-inflammatory mediator, under non-inflammatory conditions, IL-33 sustains Tregs, eosinophils, as well as ILC2 to keep an anti-inflammatory state in adipose tissue (see Figure). IL-33 is also involved in the formation of brown adipocytes from adipocyte precursors by a mechanism involving IL-13 and the endogenous opioid Met-Enkephalin secreted by activating ILC2s [2]. A direct negative role of IL-33 on adipocyte differentiation has been reported recently [3].

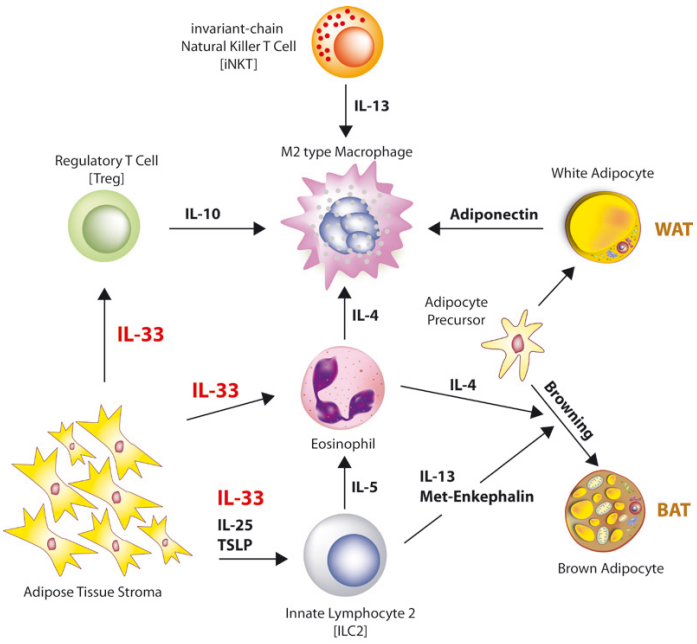


FIGURE: Control of adipose tissue homeostasis.

LIT: [1] The "Big Bang" in obese fat: Events initiating obesity-induced adipose tissue inflammation: F.M. Wensveen, et al.; Eur. J. Immunol. **45,** 2446 (2015) • **[2]** Activated type 2 innate lymphoid cells regulate beige fat biogenesis: M.W. Lee, et al.; Cell **160,** 74 (2015) • **[3]** Regulation of de novo adipocyte differentiation through crosstalk between adipocytes and pre-adipocytes: T.D. Challa, et al.; Diabetes **64,** 4075 (2015)

UNIQUE

NEW Highly Active Human IL-33 Proteins

IL-33 (oxidation resistant) (human) (rec.)

AG-40B-0160	Untagged	10 µg | 100 µg
AG-40B-0167	His-Tag	10 µg | 100 µg

LIT: Oxidation of the alarmin IL-33 regulates ST2-dependent inflammation: E.S. Cohen, et al.; Nat. Commun. **6,** ID8327 (2015)

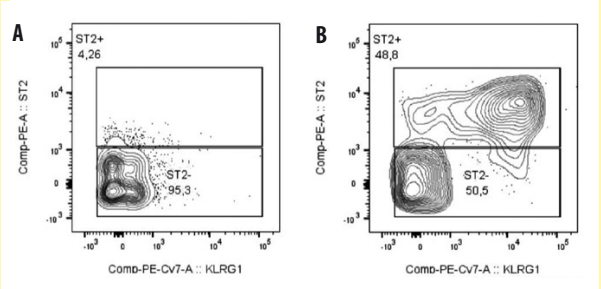


FIGURE: Activation *in vivo* of Innate Lymphoid Cells 2 (ILC2) by IL-33 (oxidation resistant) (human) (rec.) (untagged) (AG-40B-0160). Method: C57BL/6 mice were injected daily for 3 days with PBS (Figure A) or IL-33 (oxidation resistant) (human) (rec.) (untagged) (AG-40B-0160) (at 0.4µg per mouse) (Figure B). At day 4, cells from bone marrows were stained and analyzed by flow cytometry. Levels of ST2 and KLRG1 on Innate Lymphoid Cells (gated as lineage negative, CD127 positive cells) are shown. *Picture courtesy of Dr G.Verdeil / Dr S. Trabanelli (Camilla Jandus Group, Department of Fundamental Oncology, University of Lausanne).*

UNIQUE

Antibody inhibiting the binding of mouse IL-33 to ST2/IL-1RAcP

IL-33 (mouse), mAb (rec.) (blocking) (Bondy-1-1)

AG-27B-0013		100 µg
AG-27B-0013PF	Preservative Free	100 µg | 500 µg | 1mg

LIT: Regulation of de novo adipocyte differentiation through crosstalk between adipocytes and pre-adipocytes: T.D. Challa, et al.; Diabetes **64,** 4075 (2015)

ST2 Antibody for Flow Cytometry

anti-ST2 (human), pAb

AG-25A-0058		100 µg
AG-25A-0058YTD	ATTO488	100 tests
AG-25A-0058YTS	ATTO647N	100 tests

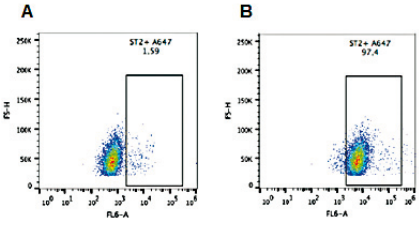


FIGURE: Detection of endogenous human ST2 with anti-ST2 (human), pAb (AG-25A-0058).

METHOD: THP1 cells were stained with anti-ST2 (human), pAb (1:100 in PBS + 2% FCS) (Figure B) or with the secondary antibody alone (Figure A) for 1h at 4°C.

Obesity & Angiogenesis

Adipose tissue is the most dynamic and plastic organ in adults. Upon exposure to different metabolic challenges, adipose tissue has the capacity to either expand or shrink according to the nutrient status. Elasticity of adipose tissue is tightly related with angiogenesis, the growth of new blood vessels, and angiogenesis plays an essential role in the modulation of adipogenesis and obesity. In growing adipose tissue, the new blood vessels contribute to adipogenesis by performing multiple functions, such as providing nutrients and oxygen to nourish adipocytes, removing waste products from the adipose tissue, carrying monocytes and neutrophils that can affect adipocyte function and also providing adipose precursors and stem cells [1].

There exist several pro-angiogenic factors secreted by adipocytes, such as leptin, adiponectin, vascular endothelial growth factor-A (VEGF-A), VEGF-B and angiopoietins (mainly ANG-1 and ANG-2) that function by stimulating proliferation and migration of endothelial cells. A recent study [2] demonstrates that angiopoietin-2 (ANG-2) overexpression induces a pro-angiogenic program in white adipose tissue (WAT), protecting against high fat diet (HFD)-induced metabolic challenges. Decreasing the angiopoietin-2 levels using a neutralization antibody (anti-Angiopoietin-2, mAb (rec.) (blocking) (Angy-2-1) (AG-27B-0016PF)) confirms the beneficial effects of endogenous ANG-2. Mechanistically, increasing vascular function and decreasing adipose tissue inflammation contribute to the beneficial effects of ANG-2. Due to the essential role of angiogenesis in the modulation of adipogenesis and obesity, anti-angiogenesis therapy has emerged as a potential treatment for obesity.

LIT: [1] The lymphatic vasculature: its role in adipose metabolism and obesity: N. Escobedo & G. Oliver; Cell Metab. **26,** S1550 (2017) • **[2]** Angiopoietin-2 in white adipose tissue improves metabolic homeostasis through enhanced angiogenesis: Y.A. An, et al.; Elife **29,** 6 (2017)

NEW

Potent ANG-2 Blocking Antibodies

anti-Angiopoietin-2, mAb (rec.) (blocking) (Angy-2-1) (preservative free)

AG-27B-0016PF — 100 µg | 500 µg | 1mg

anti-Angiopoietin-2 (human), mAb (rec.) (blocking) (Angy-1-4) (preservative free)

AG-27B-0015PF — 100 µg | 500 µg | 1mg

Also Available:

Angiopoietin-2 (human) (rec.) AG-40B-0114

Angiopoietin-2 (mouse) (rec.) AG-40B-0131

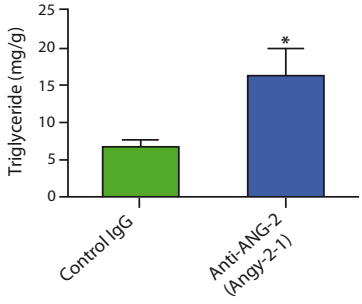


FIGURE: Antagonizing Angiopoietin-2 *in vivo* with anti-ANG-2, mAb (rec.) (blocking) (Angy-2-1) (AG-27B-0016PF) increases triglyceride levels.

METHOD: After High Fat Diet (HFD) challenges for five weeks in wild-type C57BL/6 mice, control IgG (left panel) or anti-ANG-2 (Clone Angy-2-1) blocking antibody (right panel) (4 µg/g body weight; twice/week) were administrated and afterwards the mice underwent metabolic analyses of the triglycerides levels from both groups.

Vascular Endothelial Growth Factor [VEGF]-related Reagents

PROTEINS	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
VEGF 164 (mouse) (rec.)	AG-40T-0044	5 µg | 20 µg	Sf9 cells	n.d.	Ms
VEGF 165 (human) (rec.)	AG-40T-0043	5 µg | 20 µg	E. coli	n.d.	Hu
VEGF 165 (human) (rec.)	AG-40T-0045	5 µg | 20 µg	Sf9 cells	n.d.	Hu
VEGFR-1, Soluble (human) (rec.)	AG-40T-0049	5 µg | 20 µg	Sf9 cells	n.d.	Hu
ANTIBODIES	**PID**	**SIZE**	**ISOTYPE**	**APPLICATION**	**SPECIES**
VEGF-A (human), mAb (3(6D3))	AG-20T-0105	200 µg	Mouse IgG1	ELISA, WB, FUNC	Hu
VEGFR-1 (human), mAb (EWC)	AG-20T-0106	100 µg	Mouse IgG1	ELISA, WB	Hu
VEGFR-1 (human), mAb (EWF)	AG-20T-0107	100 µg	Mouse IgG1	ELISA, IP, WB	Hu

A Complete Panel of Angiogenesis-related Reagents is available on www.adipogen.com

APPLICATIONS: FUNC: Functional Application; **ICC:** Immunocytochemistry; **IHC:** Immunohistochemistry; **IP:** Immunoprecipitation; **WB:** Western blot
FORMULATION: PF = Preservative free **SPECIES: Dg** = Dog; **Hu** = Human; **Mo** = Monkey; **Ms** = Mouse; **Rt** = Rat; **Rb** = Rabbit

Factors that Regulate WAT Browning and Thermogenesis

Brown adipose tissue (BAT) found in hibernating animals, also exists in human around the neck and collarbone. Brown adipose tissue (BAT) is the main site of adaptive thermogenesis, using a specific brown fat protein, uncoupling protein 1 (UCP1) that dissipates the mitochondrial membrane potential energy as heat instead of producing ATP. The ability of BAT to protect against obesity and metabolic diseases has traditionally been attributed to its capacity to utilize glucose and lipids for thermogenesis. However, BAT might also have a secretory role, which could contribute to the systemic consequences of BAT activity. Several BAT-derived molecules (**called Batokines**) acting in a paracrine, autocrine or endocrine manner have been identified. These Batokines control expansion and activity of BAT and the extent of browning of white adipose tissue. They also promote hypertrophy and hyperplasia of BAT, vascularization, innervation and blood flow, processes that are all associated with BAT recruitment when thermogenic activity is enhanced. Some Batokines also target peripheral tissues such as liver, pancreas, white adipose tissue, bone and heart and bone, and affect systemic metabolism by interacting with the central nervous system (CNS).

REVIEW: Brown adipose tissue as a secretory organ. F. Villarroya, et al.; Nat. Rev. Endocrinol. **13,** 26 (2017)

Overview of Important Batokines:

- **Fibroblast Growth Factor 21 (FGF-21)** is induced in BAT by cold exposure and induces the thermogenic program in brown adipocytes by interacting with FGF receptor/β-Klotho. FGF-21 is also expressed in organs such as liver or skeletal muscle. Metabolic benefits of FGF-21 include weight loss, glucose and lipid metabolism and insulin sensitivity. FGF-21 also acts directly in the brain.
- **Interleukin-6 (IL-6),** released by skeletal muscle and by BAT in response to exercise, promotes insulin sensitivity, is required for the induction of browning of WAT and acts on the pancreas and the brain (see page 10).
- **Nrg4 (Neuregulin-4)** belongs to a small family of EGF-like (EGFL) domain-containing proteins that are synthesized as transmembrane precursor and undergo proteolytic cleavage. Nrg4 is a cold-induced adipokine, highly expressed in adipose tissue, enriched in brown fat. It promotes neurite outgrowth and protects against diet-induced insulin resistance and hepatic steatosis through attenuating hepatic lipogenic signaling.
- **CTHRC1 (Collagen Triple Helix Repeat Containing 1)** is expressed in BAT but its role is still unclear (see page 5).
- **Soluble form of the LDL Receptor (sLR11)** suppresses thermogenesis in brown adipocytes, despite being increased by cold-induced activation in BAT.
- **Angiopoietin-like 8 (ANGPTL8 or Betatrophin)** is induced in BAT in response to cold but the biological significance of enhanced ANGPTL8 release by activated BAT is unclear. ANGPTL8 can repress the activity of lipoprotein lipase.
- **BMPs (Bone Morphogenetic Protein)** promote brown fat formation and act on the central nervous system to regulate thermogenesis.
- **VEGF-A and VEGF-B (Vascular Endothelial Growth Factor A and B)** regulate angiogenesis, thermogenesis and macrophage function (see page 7).
- **Slit2-C** activates a thermogenic PKA pathway in adipocytes.
- **Lipocalin Prostaglandin D Synthase (LPGDS)** synthesizes D-series prostaglandins. It is highly regulated in BAT and plays a role in lipid and carbohydrate utilization.
- **Adenosine** is released from BAT during stimulation of sympathetic nerves and activates a thermogenic program. Adenosine protects mice from diet-induced obesity.
- **Endocannabinoid system** and metabolites, such as **FFA (Free Fatty Acid), Retinaldehyde, Retinoic Acid** and **Lactate** are released from BAT and play a role in thermogenic activation.

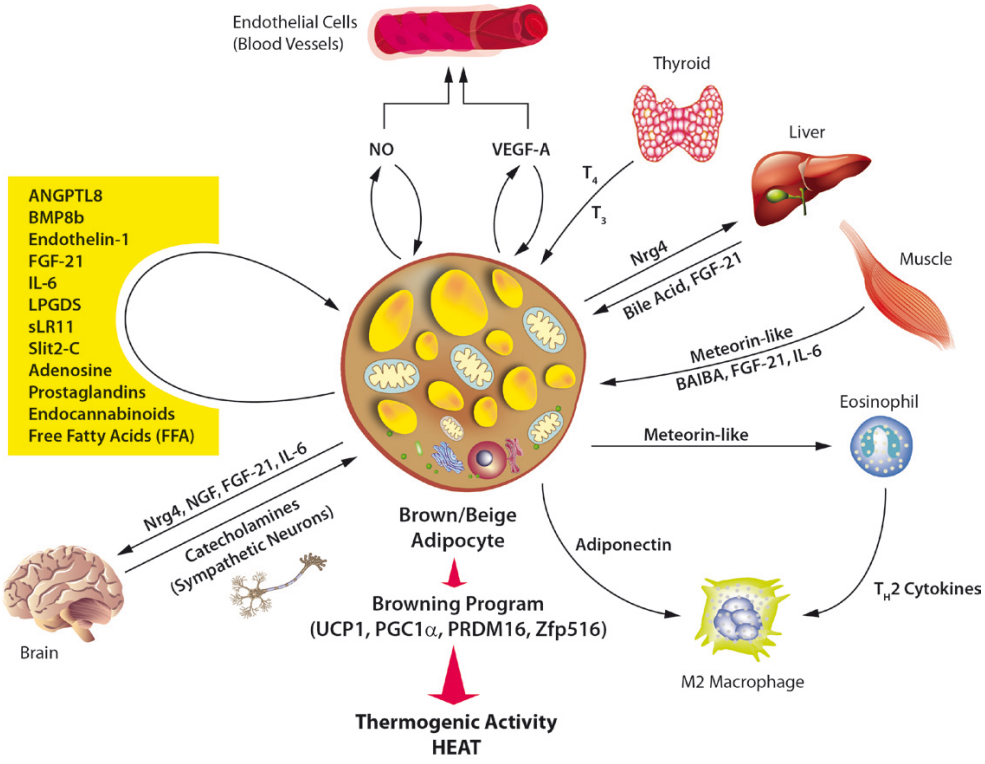


FIGURE: The autocrine and paracrine factors released by brown adipocytes.

*Adapted from F. Villarroya, et al.; Nat. Rev. Endocrinol. **13,** 26 (2017)*

Browning Inducers not expressed by BAT:

- **Cold exposure** is a strong inducer of brown cells. Thermogenic activity is regulated by a canonical β-adrenergic receptor pathway via the sympathetic nervous system. The **TRPM8 channel** is a cold-sensing cation channel present in sensing neurons that has a role in detecting environmental temperature.
- **Catecholamines** activate β-adrenergic receptors at the surface of brown adipocytes and increase the intracellular cAMP level to activate the termogenic program.
- **PPARs** are master regulators of adipogenesis. Recently, PPAR-γ activators thiazolidinediones were shown to promote WAT browning as well (see page 11).
- Cold-induced conversion of cholesterol to **Bile acid** shapes the gut microbiome and promotes adaptive thermogenesis.
- The neuropeptide **Orexin and its Receptors** are also involved in the induction of browning and affect brown fat thermogenesis.
- **Meteorin-like Protein** is a novel adipokine expressed by adipose tissue being downregulated upon caloric restriction. Meteorin-like is secreted by muscles during exercise and converts white adipose cells into brown fat tissue. This activation of fat browning is the consequence of a direct effect of meteorin-like on eosinophils in WAT that secretes IL-4 and IL-13, which promotes the activation of adipose tissue macrophages as well as the thermogenic program.
- **T3** (Triiodothyronine) and T4 exert effects locally to promote thermogenesis.
- **3-Aminoisobutyric acid (BAIBA)** is a browning molecule secreted from contracting muscles.

Protein Modulators & Inducers of Brown Adipose Tissue (BAT)

RECOMBINANT PROTEINS	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
Betatrophin (human):Fc (human) (rec.)	AG-40B-0145	10 µg | 3 x 10 µg	HEK 293 cells	<0.1EU/µg	Hu
Betatrophin (mouse) (rec.)	AG-40B-0144	10 µg | 3 x 10 µg	CHO cells	<0.1EU/µg	Ms
Betatrophin (mouse):Fc (human) (rec.)	AG-40B-0142	10 µg | 3 x 10 µg	HEK 293 cells	<0.1EU/µg	Ms
FGF-21 (human) (rec.)	AG-40A-0091	10 µg | 50 µg	HEK 293 cells	<0.1EU/µg	Hu
FGF-21 (human):Fc (human) (rec.)	AG-40A-0095	10 µg | 50 µg	HEK 293 cells	<0.1EU/µg	Hu
FGF-21 (mouse) (rec.)	AG-40B-0143	10 µg | 3 x 10 µg	HEK 293 cells	<0.01EU/µg	Ms
FGF-21 (mouse) (rec.)	CHI-MF-102FGF21	10 µg | 50 µg	HEK 293 cells	<0.06EU/µg	Ms
FGF-21 (mouse):Fc (human) (rec.)	AG-40A-0097	10 µg | 50 µg	HEK 293 cells	<0.1EU/µg	Ms
NEW **Meteorin-like (mouse) (rec.)**	AG-40B-0149	10 µg | 3 x 10 µg	HEK 293 cells	<0.1EU/µg	Ms
NEW **Neuregulin-4 (human) (rec.)**	AG-40B-0155	10 µg | 3 x 10 µg	E. coli	<0.01EU/µg	Hu
NEW **Neuregulin-4 (mouse) (rec.)**	AG-40B-0159	10 µg | 3 x 10 µg	E. coli	<0.01EU/µg	Hu, Ms
NEW **Slit2 (C fragment) (human) (rec.)**	AG-40B-0168	10 µg | 3 x 10 µg	HEK 293 cells	<0.01EU/µg	Hu

NEW

UCP1-dependent Thermogenesis Inducer through CK2

Inhibition of Casein Kinase 2 (CK2) promotes brown adipocyte biogenesis, leads to an increase in whole-body energy expenditure and ameliorates diet-induced obesity and insulin resistance in mice *in vivo* by promoting UCP1-dependent thermogenesis.

LIT: Phosphoproteomics identifies CK2 as a negative regulator of beige adipocyte thermogenesis and energy expenditure: K. Shinoda, et al.; Cell Metab. **22,** 997 (2015)

CK2 Inhibitor 10
AG-CR1-3626 — 1 mg | 5 mg

CX-4945 . HCl
AG-CR1-3629 — 1 mg | 5 mg | 25 mg

DMAT
AG-CR1-3654 — 1 mg | 5 mg | 25 mg

INDY
AG-CR1-3665 — 5 mg | 25 mg

LATEST INSIGHT

Various Browning Inducers – Described in Literature

3-Aminoisobutyric acid (Contraction-induced myokine)
AG-CR1-3596 — 250 mg | 1 g

Harmine (UCP1-dependent thermogenesis inducer)
AG-CN2-0510 — 10 mg | 50 mg | 250 mg

Miglitol (α-Glucosidase inhibitor)
AG-CR1-3635 — 10 mg | 50 mg

Papaverine . HCl (PDE10A inhibitor)
AG-CN2-0414 — 1 g | 5 g

PF-2545920 (PDE10A inhibitor)
AG-CR1-3636 — 1 mg | 5 mg | 25 mg

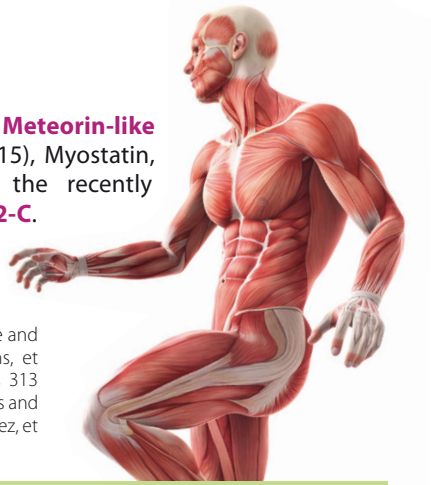
Rutin . trihydrate (Brown fat activator)
AG-CN2-0408 — 5 g

YM-254890 (Potent and selective Gαq family inhibitor)
AG-CN2-0509 — 500 µg | 1 mg

APPLICATIONS: **FUNC:** Functional Application; **ICC:** Immunocytochemistry; **IHC:** Immunohistochemistry; **IP:** Immunoprecipitation; **WB:** Western blot
FORMULATION: **PF** = Preservative free **SPECIES:** **Dg** = Dog; **Hu** = Human; **Mo** = Monkey; **Ms** = Mouse; **Rt** = Rat; **Rb** = Rabbit

Myokines: Muscle, Exercise & Obesity

Exercise training enhances muscular endurance and strength, expends calories, exerts beneficial effects on systemic metabolism and combats the development of common diseases such as obesity and type 2 diabetes, by adaptive structural and metabolic changes in skeletal muscle, including a change in the type of muscle fiber, mitochondrial biogenesis and angiogenesis. Additionally, skeletal muscles secrete cytokines and growth factors, called myokines that can potentially act in an autocrine, a paracrine and/or an endocrine manner to modulate metabolic, inflammatory and other processes. Several contraction-regulated myokines have been described including **ANGPTL4,** Apelin, BDNF, **FGF-21,** FSTL1, **IL-6, IL-7, IL-8, IL-15, Irisin, LIF,** MCP-1, **Meteorin-like protein**, Myonectin (CTRP15), Myostatin, PAI-1, **PEDF, VEGF** and the recently described **Asprosin** or **Slit2-C**.



SELECTED REVIEWS:

Exercise and regulation of adipokine and myokine production: S.W. Goergens, et al.; Prog. Mol. Biol. Transl. Sci. **135,** 313 (2015) • Crosstalk between adipokines and myokines in fat browning: A. Rodríguez, et al.; Acta Physiol. **219,** 362 (2017)

Myokine: Protein or metabolite that is produced and secreted by muscle fibers and exerts either paracrine or endocrine effects.

NEW

Asprosin

Asprosin is a new fasting-induced protein hormone that targets the liver to increase plasma glucose levels. Asprosin is the C-terminal cleavage product of the protein Fibrillin-1. Asprosin is secreted from white adipose tissue and increases hepatic glucose production by using cAMP as a second messenger, leading to activation of protein kinase A in the liver. Reduction of asprosin levels protect against metabolic syndrome-associated hyperinsulinism.

ELISA KITS	COMING SOON	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Asprosin (human) Matched Pair Detection Set		AG-46B-0012	5 x 96 wells	100 pg/ml	0.156 to 10 ng/ml	C, S
RECOMBINANT PROTEIN	**BULK AVAILABLE**	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
Asprosin (human) (rec.) (His)		AG-40B-0174	10 µg | 3 x 10 µg	E. coli	<0.01EU/µg	Hu
Asprosin (human) (rec.) (His)		AG-40B-0174T	100 µg	E. coli	<0.1EU/µg	Hu
ANTIBODIES		PID	SIZE	ISOTYPE/SOURCE	APPLICATION	SPECIES
anti-Asprosin, mAb (Birdy-1)		AG-20B-0073	100 µg	Mouse IgG1	WB	Hu, Ms
anti-Asprosin (human), mAb (Birdy-2)		AG-20B-0074	100 µg	Mouse IgG2a	WB	Hu

Selected Myokines: Interleukin-6 and Irisin

Several cytokines including IL-6, IL-7, IL-8, IL-15, LIF and MCP-1 have been shown to be secreted from muscle after endurance. IL-6 is the best characterized myokine implicated as a co-inducer of the development of obesity-associated insulin resistance, which precedes the development of type 2 diabetes (T2D). The role of irisin is still under debate. Initially, described as a browning inducer, recent studies suggest an involvement in cortical bone mass, β-cell proliferation and insulin secretion.

LIT: From cytokine to myokine: the emerging role of interleukin-6 in metabolic regulation: M. Pal, et al.; Immunol. Cell Biol. **92,** 331 (2014) (Review) • The myokine irisin is released in response to saturated fatty acids and promotes pancreatic beta-cell survival and insulin secretion: A. Natalicchio, et al.; Diabetes **(Epub ahead of print)** (2017)

PROTEINS	PID	SIZE	SOURCE	ENDOTOXIN	SPECIES
IL-6 (human) (rec.) (His)	CHI-HF-20106	10 µg | 50 µg	HEK 293 cells	<0.01EU/µg	Hu
IL-6 (human):Fc (human) (rec.)	CHI-HF-21006	50 µg | 3 x 50 µg	CHO cells	<0.06EU/µg	Hu
IL-6 (mouse):Fc (human) (rec.)	AG-40B-0108	10 µg | 3 x 10 µg	HEK 293 cells	<0.01EU/µg	Ms
Irisin (rec.) (CHO)	AG-40B-0136	10 µg | 3 x 10 µg	CHO cells	<0.01EU/µg	Hu, Ms
Irisin (rec.) (E. coli)	AG-40B-0103	10 µg | 5 x 10 µg	E. coli	<0.1EU/µg	Hu, Ms
ELISA KITS	PID	SIZE	SENSITIVITY	RANGE	SAMPLES
Cymax IL-6 (human) ELISA Kit	YIF-LF-EK0260	96 wells	1.160 pg/ml	4.68 to 300 pg/ml	C, P, S, L
Cymax IL-6 (mouse) ELISA Kit	YIF-LF-EK0270	96 wells	1.138 pg/ml	7.8 to 500 pg/ml	C, P, S, L
Cymax IL-6 (rat) ELISA Kit	YIF-LF-EK0224	96 wells	26.643 pg/ml	62.5 to 4000 pg/ml	C, P, S, L
Irisin Competitive ELISA Kit	AG-45A-0046Y	96 wells | 2 x 96 wells	1 ng/ml	0.001 to 5 µg/ml	C, P, S

SAMPLES: C: Cell Culture Supernatant, **P:** Plasma, **S:** Serum; **U:** Urine; **L:** Cell Lysate; **F:** CSF
APPLICATIONS: FUNC: Functional Application; **ICC:** Immunocytochemistry; **IHC:** Immunohistochemistry; **IP:** Immunoprecipitation; **WB:** Western blot
FORMULATION: PF = Preservative free
SPECIES: Dg = Dog; **Hu** = Human; **Mo** = Monkey; **Ms** = Mouse; **Rt** = Rat; **Rb** = Rabbit

AMPK Modulators

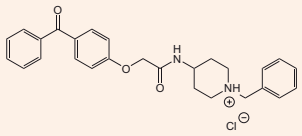
AMPK (AMP-activated protein kinase) plays a role in cellular energy homeostasis, regulating several intracellular systems including hepatic fatty acid oxidation and ketogenesis, inhibition of cholesterol synthesis, lipogenesis and triglyceride synthesis, stimulation of skeletal muscle fatty acid oxidation and muscle glucose uptake as well as modulation of insulin secretion by pancreatic β cells.

SELECTED REVIEW ARTICLE: Past strategies and future directions for identifying AMP-activated protein kinase (AMPK) modulators: S.E. Sinnett & J.E. Brenman; Pharmacol. Ther. **143,** 111 (2014)

BULK

AdipoRon . HCl (water soluble) **Original Source**
AG-CR1-0156 10 mg | 50 mg

Formula: $C_{27}H_{28}N_2O_3$. HCl . H_2O
MW: 428.5 . 36.5 . 18.0
CAS: 924416-43-3 (free base)



AdipoR agonist. AMPK & PGC1α activator. Improves diabetes.

AICAR (Potent AMPK activator) BULK
AG-CR1-0061 10 mg | 50 mg | 100 mg

AdipoRon (AMPK & PGC1α activator) BULK
AG-CR1-0154 10 mg | 50 mg

Compound 112254 . HCl (water soluble) **(AMPK activator)**
AG-CR1-0157 10 mg | 50 mg

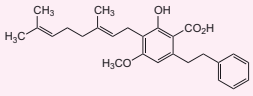
MOTS-c (human) (AMPK inducer) NEW
AG-CP3-0026 1 mg | 5 mg

PPAR (Peroxisome Proliferator-activated Receptor) Agonists

NEW

Amorfrutin B
AG-CN2-0464 500 µg

Formula: $C_{26}H_{32}O_4$
MW: 408.5
CAS: 1174387-94-0
Source: *Amorpha fruticosa*



Natural PPARγ agonist with potent glucose-lowering properties.

Also available: **Amorfrutin A (AG-CN2-0462)**

Astaxanthin (PPARα agonist & PPARγ antagonist) BULK
AG-CN2-0055 5 mg | 25 mg

Ciglitazone (Selective PPARγ agonist) BULK
AG-CR1-0033 1 mg | 5 mg | 25 mg

GW1929 (Selective PPARγ agonist) BULK
AG-CR1-0116 1 mg | 5 mg | 25 mg

GW501516 (Potent and selective PPARδ agonist) NEW
AG-CR1-3641 1 mg | 5 mg | 25 mg

Ionomycin (free acid) (PPARγ ligand with a unique binding mode)
AG-CN2-0416 1 mg | 5 mg

Pioglitazone (Selective PPARγ agonist) BULK
AG-CR1-0067 1 mg | 5 mg | 25 mg

Rosiglitazone . maleate (Potent PPARγ agonist) BULK
AG-CR1-3571 25 mg | 100 mg | 1 g

Pseudolaric acid B (PPARα agonist)
AG-CN2-0083 100 µg | 1 mg

Troglitazone (Potent and selective PPARγ agonist)
AG-CR1-3565 5 mg | 5 mg | 25 mg

WY-14643 [Pirinixic acid] (Potent PPARα activator)
AG-CR1-3566 10 mg | 50 mg | 250 mg

LATEST INSIGHT

Leptin Sensitizers

Celastrol
AG-CN2-0460 5 mg | 10 mg | 50 mg

Withaferin A
AG-CN2-0490 1 mg | 5 mg | 10 mg

LATEST INSIGHT

Immunometabolism Modulators

Shikonin (PKM2/Glycolysis inhibitor)
AG-CN2-0487 10 mg | 50 mg

N-Acetyl-D-glucosamine (Hexokinase inhibitor)
AG-CN2-0489 250 mg | 1 g | 5 g

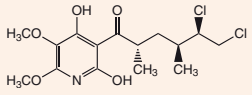
Selection of a Broad Range of Metabolic Research Reagents

BULK

Atpenin A5 (synthetic) — Original Source

AG-CN2-0100 — 250 µg | 1 mg

Formula: $C_{15}H_{21}Cl_2NO_5$
MW: 366.2
CAS: 119509-24-9



Potent and specific mitochondrial complex II (succinate-ubiquinoneoxidoreductase) inhibitor.

AK-7 (Brain-permeable SIRT2 inhibitor)
AG-CR1-3511 — 5 mg | 25 mg

Amlexanox (Selective TBK1 and IKKε inhibitor)
AG-CR1-3579 — 10 mg | 50 mg

BMS-309403 (Potent and selective FABP4 inhibitor)
AG-CR1-3640 — 1 mg | 5 mg | 25 mg

EM574 (Orexigenic; Motilin receptor agonist)
AG-CN2-0102 — 250 µg | 1 mg

Emodin (Potent selective 11β-HSD1 inhibitor)
AG-CN2-0457 — 50 mg | 250 mg

Empagliflozin (SGLT-2 inhibitor)
AG-CR1-3619 — 10 mg | 50 mg

Genipin (UCP2 inhibitor)
AG-CN2-0481 — 25 mg | 100 mg | 500 mg

Glyburide (USP) (Antidiabetic)
AG-CR1-3613 — 1 g | 5 g | 10 g

Isoliquiritigenin (Antidiabetic/Antihyperglycemic)
AG-CN2-0459 — 10 mg | 50 mg

Kaempferitrin (Insulinomimetic/Hypoglycemic)
AG-CN2-0039 — 1 mg | 5 mg

Linagliptin (DPP4 inhibitor)
AG-CR1-3618 — 10 mg | 50 mg

Orlistat (DAGLα inhibitor/Antiobesity) BULK
AG-CN2-0050 — 50 mg | 250 mg

Pellitorine (α-Glucosidase inhibitor) BULK
AG-CN2-0009 — 1 mg | 5 mg | 25 mg

Resveratrol (Potent SIRT1 activator) BULK
AG-CN2-0033 — 50 mg | 100 mg | 500 mg

Salsalate (Anti-inflammatory/Antidiabetic)
AG-CR1-3574 — 1 g | 5 g

Sirtinol (Cell permeable SIRT1 inhibitor) BULK
AG-CR1-0055 — 1 mg | 5 mg | 25 mg

Skyrin (Antidiabetic)
AG-CN2-0001 — 1 mg

Stevioside (Antidiabetic)
AG-CN2-0077 — 10 mg | 50 mg

Suramin . 6Na (SIRT1 & SIRT5 inhibitor)
AG-CR1-3575 — 50 mg | 250 mg

(±)-Verapamil . HCl (USP Grade) (Antidiabetic)
AG-CR1-3627 — 100 mg | 1 g | 5 g

Vitexin (α-Glucosidase inhibitor)
AG-CN2-0425 — 5 mg | 25 mg

LATEST INSIGHT

Microbiota-related Reagents

Indole-3-carbinol NEW
AG-CR1-3637 — 500 mg | 5 g
Aryl hydrocarbon receptor (AhR) agonist. Used to increase the population of innate lymphoid cells 3 (ILC3) in gut microbiota.

***trans*-Indole-3-acrylic acid** NEW
AG-CR1-3677 — 250 mg | 1 g
Tryptophan metabolite produced by human microbiota (intestinal commensale *Peptostreptococcus sp.*). Involved in keeping the intestinal barrier intact.

LATEST INSIGHT

NEW Obesity Research Tools

GW311616A . HCl NEW
AG-CR1-3632 — 1 mg | 5 mg | 25 mg
Potent, selective intracellular neutrophil elastase inhibitor. Reverses insulin resistance and body weight gain in HFD-fed mice.

Niclosamide NEW
AG-CR1-3643 — 100 mg | 1 g

Niclosamide . ethanolamine NEW
AG-CR1-3644 — 25 mg
Positive allosteric neuropetide Y4 receptor ligand that increases energy expenditure and lipid metabolism through mitochondrial uncoupling.

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