Hormone Sensitive Lipase

Contents

- 1 Overview
 - ◆ 1.1 Hormone Sensitive Lipase (HSL)
 - ↑ 1.2 HSL, a drug target for Diabetes and Metabolic Disease
 ↑ 1.3 Type 2 Diabetes

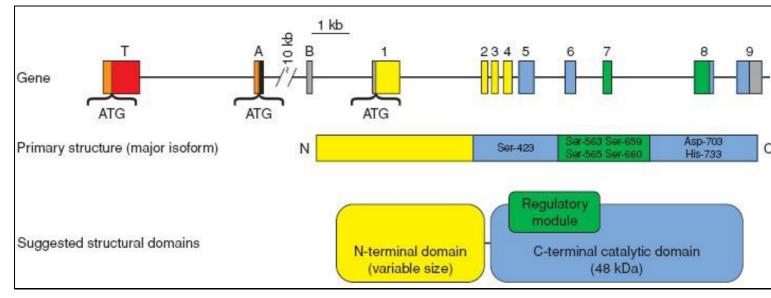
 - ◆ 1.4 Metabolic Disease
- 2 Non-patent Literature Search and Analysis
 - ◆ 2.1 Search Strategy
 - ◆ 2.2 Scientific papers on chemical compounds inhibiting or stimulating HSL
 ◆ 2.3 General scientific studies on HSL found from STN
- 3 Patentability Search
 - ◆ 3.1 Sequence Search and Analysis
 - 3.2 Sequence Search Stratagy
- 4 Like this report?
 - 4.1 Micropatent Search and Analysis
 - ◊ 4.1.1 Final Patent list (with bibliographic information)
 - ♦ 4.1.2 Overall IP activity
 - ♦ 4.1.3 Key Players (those with more than 2 patents) ♦ 4.1.4 Other players (those with 1 patent)
- 5 Patent Analysis
 - ◆ 5.1 Patent Distribution (Sample analysis of 25 records only)
- 6 Dashboard (Sample based on 25 analyzed records)
- 7 Interactive Mindmap
- 8 Like this report?
- 9 Contact Dolcera

Overview

Hormone Sensitive Lipase (HSL)

- Hormone-sensitive lipase is an enzyme expressed in multiple tissues. It plays number of roles in lipid metabolism.
- Major isoform of HSL gene is a single polypeptide with molecular mass of approximately 84kDa, its has 3 major domains catalytic, regulatory domain congaing several plosphorylation sistes, and n-terminal domain involved in protein-protein and protein-lipid interactions.

 • Altererd expression of HSL in different celltypes may be associated with several pathological states, including obesity, atherosclerosis, Type II
- Diabetes.
- · HSL has lipolytic activity against not only triacylglycerol, but also against diacylglycerol, monoacylglycerol, and cholesteryl esters.
- · Gene and protein structure of the HSL given below.





Gene: exon/intron organization of the human HSL gene. Exons T and A encode 300 and 43 additional amino acids respectively, located at the N-terminus.

Protein: Linear representation of the amino acid sequence of adipocyte HSL (numbering of the rat sequence).

Domains: N-terminal binding domain, a C-terminal catalytic domain, harbouring the catalytic triad and a regulatory module containing multiple phosphorylation sites. Source

• Fates of HSL products in different cell types.

Tissue	Major product	Function
Adipose tissue	Fatty acids	Export for oxidation
Muscle, heart	Fatty acids	Oxidation
Pancreatic β-cells	Fatty acids	Signalling?
Adrenals, testes, ovaries	Cholesterol	Substrate for steroidogenesis
Macrophages	Cholesterol	Export (via high-density lipoprotein)
Placenta, ovaries	Steroids	Transcriptional control
Mammary gland	Cholesterol	Milk component, membranogenesis

- HSL protein sequence Source1, Source2
 HSL mRNA sequence Source

HSL, a drug target for Diabetes and Metabolic Disease

A few points suggesting HSL as a drug tareget for Type 2 Diabetes and Metabolic Disease:

- There are conflicting data on whether altered expression or activity of HSL is involved in familial combined hyperlipidaemia. Similar debate surrounds the role of altered levels of HSL in obesity and in insulin resistance, two linked conditions which predispose to Type II Diabetes. Source

- Elevated levels of plasma nonesterified fatty acids are associated with insulin resistance.
 A promoter variant of HSL,?60C?G, has been implicated in a number of conditions, which exhibits a 40% decrease in promoter activity, has been associated with increased insulin sensitivity in women and decreased levels of plasma non-esterified fatty acid in men.
 Population studies suggest that a polymorphic marker in the HSL gene is in linkage disequilibrium with an allele that increases susceptibility to abdominal obesity, which is itself a risk factor for Type II diabetes. Source
 Over expression of HSL prevents lipid accumulation (from cell line studies). Source
- Over expression of HSL prevents lipid accumulation (from cell line studies). Source

Type 2 Diabetes

- Type 2 Diabetes Mellitus is a group of metabolic diseases characterized by high blood sugar (glucose) levels, which result from defects in
- insulin secretion, or insulin resistance, or both.
 Insulin, a hormone produced by the pancreas, which controls blood glucose levels.
 Insulin triggers glucose uptake by the cells. Some part of the glucose can be converted to concentrated energy sources like glycogen, triangular part fath points. triacylglycerols, or fatty acids.

Metabolic Disease

- A person with abnormal glucose tolerance (IGT or diabetes) will be found to have at least one or more of the other cardiovascular disease (CVD) risk components.

 • This clustering has been labelled variously as Syndrome X, the Insulin Resistance Syndrome, the Metabolic Syndrome or Metabolic disease.

Non-patent Literature Search and Analysis

• In non-patent literature searching we got 17 records/articles. Out of these 8 articles are talking about chemical compounds that are involved in inhibition or stimulation of hormone-sensitive lipase (HSL) and 9 articles are scientific review papers.

Search Strategy

S.No.	Database	e Key-Words		Search Queries	No. of
		Main Terms	Alternative Terms		Records
1	BIOSIS	Horrnone sensitive lipase Type 2 diabetes	Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS • Diabetes, NIDDM, IDDM,	 L4 1576 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR HSL OR LHS OR LIPE OR (HORMONESENSITIVE (A) LIPASE#) L5 266641 DIABET? OR NIDDM OR IDDM OR T2DM L6 30 L4 (S) L5 	30
2	EMBASE	 Hormone sensitive lipase Type 2 diabetes 	Lipase-normone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS, L11706, U40001, Q05469	 L2 1203 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR (HORMONESENSITIVE (W) LIPASE#) OR LIPE OR HSL OR LHS OR L11706 OR U40001 OR Q05469 L3 258095 DIABET? OR NIDDM OR IDDM OR T2DM L4 32 L2 (S) L3 	32
3	DISSABS	 Hormone sensitive lipase Type 2 diabetes 	Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS Diabetes, NIDDM, IDDM,	 L1 131 (HORMONE (W) SENSITIVE (2A) LIPASE) OR (HORMONESENSITIVE (W) LIPASE) OR LIPE OR HSL OR LHS L2 6000 DIABET? OR NIDDM OR IDDM OR T2DM L3 5 L1 (S) L2 	5
4	MEDLINE	• Hormone sensitive		• L1 1331 (HORMONE (W) SENSITIVE (2A) LIPASE) OR	29

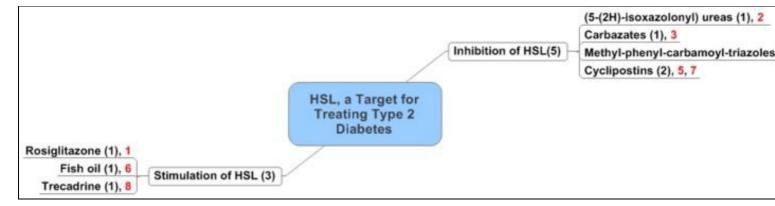
		lipase • Type 2 diabetes	Hormonesensitive lipase, LIPE, HSL, LHS, L11706, U40001, Q05469 • Diabetes, NIDDM, IDDM, T2DM	(HORMONESENSITIVE (W) LIPASE) OR LIPE OR HSL OR LHS OR L11706 OR U40001 OR Q05469 • L2 286108 DIABET? OR NIDDM OR IDDM OR T2DM • L3 29 L1 (S) L2	
5	BIOTECHNO	Hormone sensitive lipase Type 2 diabetes	 Hormone-sensitive lipase, Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS, L11706, U40001 Diabetes, NIDDM, IDDM, T2DM 	L1 398 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR (HORMONESENSITIVE (W) LIPASE#) OR LIPE OR HSL OR LHS OR L11706 OR U40001 L2 18139 DIABET? OR NIDDM OR IDDM OR T2DM L3 10 L1 (S) L2	10
6	BIOTECHABS	Hormone sensitive lipase Type 2 diabetes	 Hormone-sensitive lipase, Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS, L11706, U40001, Q05469 Diabetes, NIDDM, IDDM, T2DM 	L1 56 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR (HORMONESENSITIVE (W) LIPASE#) OR LIPE OR HSL OR LHS L2 9716 DIABET? OR NIDDM OR IDDM OR T2DM L3 5 L1 (S) L2	5
7	CONFSCI	Hormone sensitive lipase Type 2 diabetes	Hormone-sensitive lipase, Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS Diabetes, NIDDM, IDDM, T2DM	L1 90 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR (HORMONESENSITIVE (W) LIPASE#) OR LIPE OR HSL OR LHS L2 1257 DIABET? OR NIDDM OR IDDM OR T2DM L3 0 L1 (S) L2	0
8	NTIS	 Hormone sensitive lipase Type 2 diabetes 	 Hormone-sensitive lipase, Lipase-hormone sensitive, Hormonesensitive lipase, LIPE, HSL, LHS Diabetes, NIDDM, IDDM, T2DM 	L1 90 (HORMONE (W) SENSITIVE (2A) LIPASE#) OR (HORMONESENSITIVE (W) LIPASE#) OR LIPE OR HSL OR LHS L2 1257 DIABET? OR NIDDM OR IDDM OR T2DM L3 0 L1 (S) L2	0

Total number of records = 106
After removing duplicates number of records = 49
Total on-targets (after removing duplicates) = 17

Accuracy of the query = 35%

Scientific papers on chemical compounds inhibiting or stimulating HSL

- Following is the distribution of the 8 articles based on Function of the compounds.
- Numbers in brackets indicate number of records in the concerned topic.
- Highlighted (in red color) numbers refer serial number in the analysis table.



Distribution of non-patent articles

• Detailed analysis of the above records is available at the following link: Analysis Table

General scientific studies on HSL found from STN

- Off the 17 articles, 9 articles are talking about variations in the HSL gene and review papers. Variations discussed are C-60G polymorphism in the promoter of the HSL gene and a dinucleotide repeat in the gene, which are associated with Type 2 diabetes, suggesting the putative role of the HSL in Type 2 diabetes. Other articles are reviews saying HSL is a potential target for treating or curing Type 2 diabetes.
 The following hyperlink takes us to the above mentioned 9 articles: Scientific studies of HSL

Patentability Search

Sequence Search and Analysis

- Sequence search on NCBI.
 Got 16 relevant patents.
 Spread sheet
- Sequence search on DGENE.
 Got 42 relevant patents.
- Spread sheet
 Sequence search on PCTGEN.
 - Got 16 relevant patents.
 - ◆ Spread sheet

Sequence Search Stratagy

S.No.	Database	Sequence Query	Key Words (to restrict unrelated patents)	Hits
1	NCBI (Blast)	HSL protein		13
2	NCBI (Blast)	• HSL mRNA		19
3	DGENE (Blast search)	HSL protein HSL mRNA	diabetesNIDDMIDDMT2DM	61
4	PCTGEN (Blast search)	HSL protein HSL mRNA		83

- In NCBI got 16 relevat patents.
 In DGENE got 42 relevant patents.
 In PCTGEN got 16 relevant patents.

Like this report?

This is only a sample report with brief analysis Dolcera can provide a comprehensive report customized to your needs

Buy the customized report from Dolcera			
Patent Analytics Services	Market Research Services	Purchase Patent Dashboard	
Patent Landscape Services	Dolcera Processes	Industry Focus	
Patent Search Services	Patent Alerting Services	Dolcera Tools	

Micropatent Search and Analysis

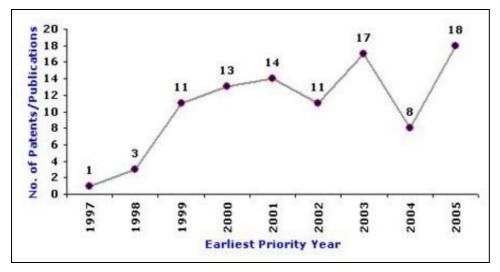
- The following is the link to search strategy on Micropat Search on micropat
 The following provides an example of how patents were analyzedSample patent analysis
- From Micropat search, 769 patents were analyzed out of which 41 are on target.

Final Patent list (with bibliographic information)

- After removing duplicates and family members of relevant patents found from NCBI, PCT Gene, DGENE and Micropat, we got 96 relevant patents.
 • Final list of patents

Overall IP activity

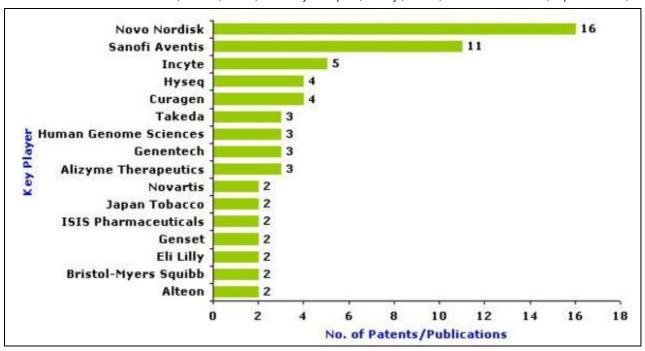
• The below graph indicates the overall patenting trend for the 96 relevant patents over time



5

Overall IP Activity
Key Players (those with more than 2 patents)

• Key players in the field (total 66 records) are Novo Nordisk, Sanofi Aventis, Incyte, Curagen, Hyseq, Alizyme Therapeutics, Genentech, Human Genome Sciences, Takeda, Alteon, Bristol-Myers Squibb, Eli Lilly ,Genset, ISIS Pharmaceuticals, Japan Tobacco, Novartis.



12

Key Players
Other players (those with 1 patent)

• In all, other players have 30 patents to their name.

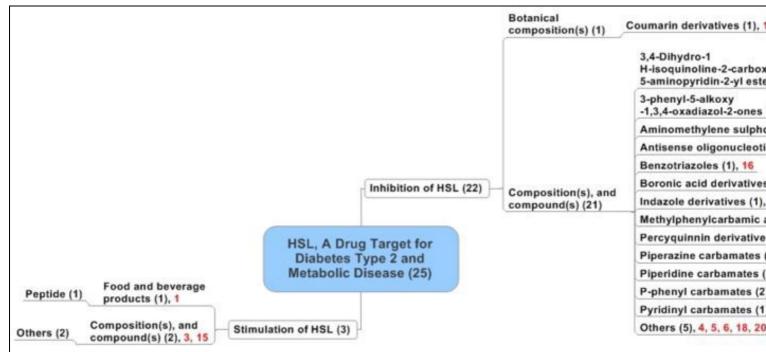
Assignee	No. of Patents/Publications	
Arena Pharmaceuticals	1	
Ares Trading	1	
Asahi Kasei Pharma Corporation	1	
Bionomics	1	
Compugen	1	
Epiontis	1	
Euroscreen	1	
Exelixis	1	
F. Hoffmann La Roche Ag	1	

Five Prime Therapeutics	1
Forschungsverbund Berlin	1
Genetics linstitute LLC, Ono Pharmaceuticals	1
GPC Biotech	1
Immunex	1
Linkagene Ltd.	1
Merck & Co	1
Metabolex	1
Micromet AG	1
Nuvelo	1
Ottawa Health Research Institute	1
President & Fellows Of Harvard College	1
Protein Design Labs	1
Regents Of The University Of California	1
Rutgers,The State University Of New Jersey	1
Schering Corporation	1
Sequenom	1
The Buck Institute For Age Research	1
The Regents Of The University Of Michigan	1
University Of Maryland Baltimore	1
Yissum Research Development Company Of The Hebrew University Of Jerusalem	1

Patent Analysis

Patent Distribution (Sample analysis of 25 records only)

- HSL refers to hormone-sensitive lipase.
- Patents distributed based on patent focus and mechanism of the action of the compounds.
- The number of patents referring to a class of compounds is given in the brackets in the figure.
- The below map is a sample representation.





Patent Distribution

• Details about patents available from the following hyperlink, analysis sheet

Dashboard (Sample based on 25 analyzed records)

- Dashboard on patents and analysis
 The given dashboard is a sample representation.

Interactive Mindmap

```
.markmap-node {
    cursor: pointer;
      .markmap-node-circle {
    fill: #fff;
             stroke-width: 1.5px;
      .markmap-node-text {
    fill: #000;
    font: 10px sans-serif;
      .markmap-link {
    fill: none;
      pre, .mw-code{
  background-color: transparent;
}
d3.xml("https://www.dolcera.com/wiki/images/Mmap22.mm", function(error, data) { if (error) throw error;
         markmap("svg#mindmap_6e83429d5abe3bf4d535fe52fe7820a7", data, {
    preset: "colorful",
    linkShape: "diagonal"
}, "xml");
});
```

Like this report?

This is only a sample report with brief analysis Dolcera can provide a comprehensive report customized to your needs

Buy the customized report from Dolcera			
Patent Analytics Services	Market Research Services	Purchase Patent Dashboard	
Patent Landscape Services	Dolcera Processes	Industry Focus	
Patent Search Services	Patent Alerting Services	Dolcera Tools	

Contact Dolcera

Samir Raiyani

Email: info@dolcera.com Phone: +1-650-269-7952