



# 2008 INVESTORS' GUIDE



Carna Biosciences, Inc. KIBC 511, 5-5-2 Minatojima-Minamimachi,







# Message from the President

Carna Biosciences, Inc. expertizes on kinases to establish drug discovery technology and aims at the development of new drugs.

#### Working as the leading kinase company

Carna Biosciences focuses on the drug discovery by targeting protein kinases, kinds of phosphotransferase enzymes. Protein kinases were originally discovered as proto-oncogenes in cancer research and it was identified 518 putative human protein kinase genes. Since the protein kinases play crucial roles in cellular signal transduction, uncontrolled kinase activities may result in diseases such as cancer and autoimmune diseases. To date, 9 kinase inhibitors have been approved for cancer therapy in the USA provided superior clinical outcome in cancer patients to compared with classical anti-cancer drugs. The launching of these kinase inhibitors called molecular-targeted drugs has greatly changed the treatment of the disease. Our primary aim is to create highly effective drugs for diseases such as cancer and autoimmune diseases. High quality kinase proteins are necessary for drug discovery research of kinase inhibitors. Carna Biosciences has generated 297 kinds (as of the end of August 2008) of high quality human recombinant kinase proteins and developed 276 kinds (as of the end of August 2008) of their kinase assays, which are used as profiling panels to investigate selectivity of kinase inhibitors. These products and services are provided to pharmaceutical companies as drug discovery services. Besides, they are utilized for internal drug discovery research at Carna Biosciences. In the future, we will provide the highly effective drugs to clinicalcare, where patients are suffering with diseases, through the products and services that we are supplying to pharmaceutical companies. We also would develop and supply the drug-candidate compounds to help the drug-discovery research of pharmaceutical companies.

#### The origin of our company name

Carna was founded in April 2003 as a bio-venture company that offers products and services necessary for kinase inhibitor drug discovery which comprises recombinant protein kinases and selectivity profiling services to pharmaceutical companies. Nowadays Carna has started a drug discovery that can generate lead compounds with internal drug discovery programs in addition to such products and services.

"Carna" is the goddess who takes care of human health, protecting the human heart and other organs as well as everyday life.Carna is a goddess of roman mythology, who takes care of human health and is said to be the root for the word "cardiac". The word "biosciences" is derived from the words "biology" and "life sciences". Carna Biosciences has created contemporary Carna goddess with protein kinase.





# **Mission**

Our mission is to develop and commercialize innovative new therapeutics which improve quality of life and lead to greater human wellness and longevity. We are strongly committed to the discovery of new drugs for the treatment of cancer and autoimmune diseases and to partnering our drug discovery technologies with global biopharmaceutical companies who share our vision to produce treatments benefiting mankind.

#### Code of conduct

- We will be thoroughly and faithfully committed and build strong relations of trust
- 2 We will always do our best to overcome difficulties
- We will respect individuality and develop 3 our imaginations and creativity



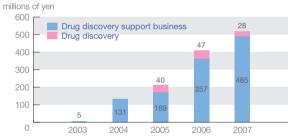
#### Selected Financial Data

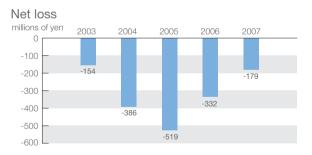
	2003	2004	2005	2006	2007
Net sales (millions of yen)	5	131	209	405	513
Ordinary loss (millions of yen)	-154	-386	-517	-269	-158
Net loss (millions of yen)	-154	-386	-519	-332	-179
Net assets (millions of yen)	329	866	358	1,652	1,435
Total assets (millions of yen)	593	1,080	564	1,938	1,622
R&D expenses (millions of yen)	94	272	392	307	256
Equity ratio (%)	55.5	80.2	63.5	85.3	88.5
Total number of issued shares (shares)	1,944	28,690	28,690	44,490	44,490

Notes: 1. The company does not issue consolidated financial stateme

 The company does not issue considered infancial statements.
Because the company was established on April 10, 2003, the FY2003 was the roughly 9-month period from April 10, 2003 to December 31, 2003.
The financial statements for the FY2005 and the FY2006 were audited by Deloitte Touche Tohmatsu based on the provisions of Article 3.7 securities listing regulations of the JASDAQ Securities Exchange, Inc. 4. The company conducted a 1-to-10 share split on May 6, 2004.

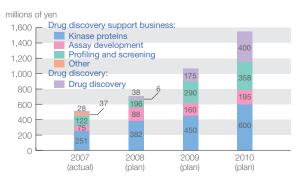
## Net sales





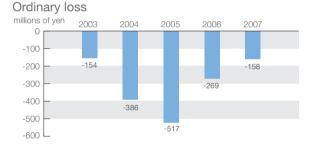
# Milestone

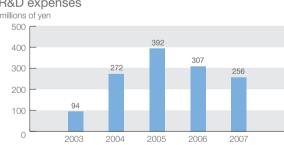
Net sales by business segment and product category - Drug discovery support businese & Drug discovery





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# R&D expenses

#### Net sales by region

- Drug discovery support business





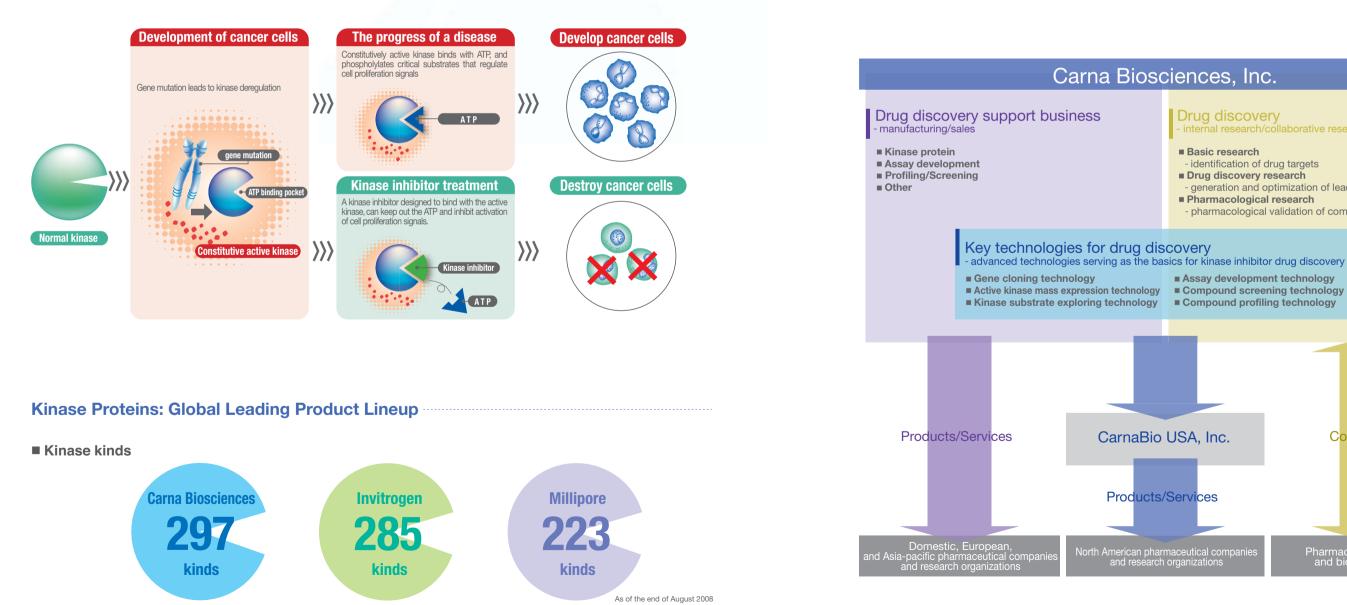
#### **Protein Kinase**

Protein kinases are enzymes that transfer a phosphate group of ATP to the specific target proteins. In human cells, a variety of extracellular stimuli generate intracellular signals of which mechanisms are reversible protein phosphorylation that are catalyzed by more than 500 members of protein kinases. The complex signaling networks by protein kinases regulate a wide range of biological events such as cell proliferation, differentiation, motility and secretion of physiologically active substances. Because of its biological importance, uncontrolled protein kinase function is known to be a cause of various diseases such as cancer, autoimmune, etc. Carna are focusing protein kinases as drug discovery targets.

#### **Kinase Inhibitors**

It has become clear that in cancer, malfunction of certain specific kinases cause abnormal proliferation and/or differentiation of cells. This finding has motivated pharmaceutical companies to pursue protein kinases as drug targets that now account for 20-30% of all drug discovery programs, and produced highly effective anti-cancer drugs with less adverse effects. Numerous kinase inhibitors are also currently undergoing preclinical and clinical evaluation.

#### Kinase inhibitor mechanism



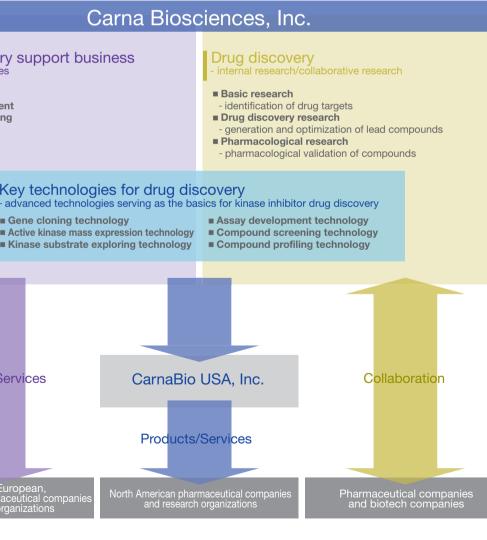
# Segment Information

Carna consists of drug discovery support business and drug discovery, that are individually operated by common policy "key technologies for kinase inhibitor drug discovery."

In the drug discovery support business, we put efforts on research and development of recombinant human kinase proteins and their assay systems, which are required for kinase inhibitor drug discovery. Our main products and services include the kinase protein, kinase assay development, selectivity profiling, and X-ray crystal structure analysis of target protein.

In the drug discovery, we have own drug discovery programs to generate lead candidates in cooperation with other pharmaceutical companies and research organizations.

# ARNA BIOSCIENCES





# Drug discovery Support business

Carna has been establishing "key drug discovery technologies" which are required for kinase inhibitor drug discovery. With these technologies, Carna offers products and services to support your research in early stage, e.g. kinase protein, kinase assay development, selectivity profiling, and X-ray crystal structure analysis of target protein. Our products and services save costs and compress time consuming early drug discovery process.

Carna supports your drug discovery providing the following products and services.

### Protein Kinase



At the end of August 2008, the products of recombinant protein comprised of 80 tyrosine kinases (including 12 active mutants), 217 serine/threonine kinases (including 2 active mutants), 7 inactive kinases and 7 inactive mutants. These active proteins are suitable for screening and profiling of kinase inhibitors. The highly purified proteins can be used for protein crystallographic study. The quantity range of the protein is from small (5 µg) to large (mg level).

Carna Biosciences will continuously develop human recombinant kinases that have physiological and pathological significance, and is aiming at offering all active kinases.

### Assay Development



Carna possesses a lot of know-how on gene cloning, protein expression and purification, substrate identification and kinase assay development. Based on this know-how, Carna has developed kinase assay against 276 kinases using ELISA, IMAP and the microfluidic technology up to August 2008. Carna's assay development products include the kinase assay kits, which are comprised of the kinase protein, its substrate, assay buffer and the protocol. Carna offers customization of the assay kits on demand.

### Profiling



Profiling has become a standard role in the kinase inhibitor drug discovery process. By profiling against the largest panel of protein kinases, a detailed selectivity is revealed. This is information supports and accelerates drug development. Carna receives compounds from customers and reports the results of profiling as a contract service. With expertise to develop kinase assays and produce high-quality enzymes, Carna offers you excellent and reliable data.

Carna's profiling service panel includes the largest, most diverse protein kinases in the world and is still under constant expansion.

As of the end of August 2008, Carna can offer 90 tyrosine kinases that include most in human genome and 185 serine/threonine kinases in the profiling panel with 1 lipid kinase. Carna has the QuickScout® select Panel (MAPK cascade panel, and pre-selected 20 tyrosine kinase and 30 serine/threonine kinase panel), of which kinases have been demonstrated to be physiologically important. The profiling data represents inhibitor potency and IC50 curves against any kinases on the profiling panel. The main platform of the profiling is microfluidic mobility shift assay using LabChip3000 (Caliper Life Sciences, Inc.).



#### Screening



Carna offers library screening services against the specific kinases as a contract service. Against 276 protein kinases, we are ready to perform high-throughput screening for your library in our facility. The main platform of library screening is the microfluidic mobility shift assay but it is also able to adapt other platforms (TR-FRET, IMAPTM, AlphaScreenTM, etc.).

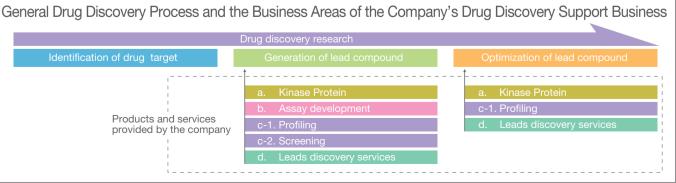
Other

#### X-ray crystal structure analysis services

Carna concluded an exclusive distributorship agreement with Crystal Genomics Inc. (South Korea, "Crystal Genomics" below) in October 2004, with the X-ray crystal structure analysis services. Carna is sole distributor of crystallographic analysis services provided by Crystal Genomics. The co-crystallization with the compound and the kinase protein could provide useful information for structure-based drug design to optimize the lead compound in the drug discovery process.

#### Leads discovery services

Carna Biosciences concluded a collaboration and services agreement with OSI Pharmaceuticals, Inc. (USA) in October 2007 for a joint venture in the cooperative development of leads discovery service focusing on kinase targets.







# Drug discovery

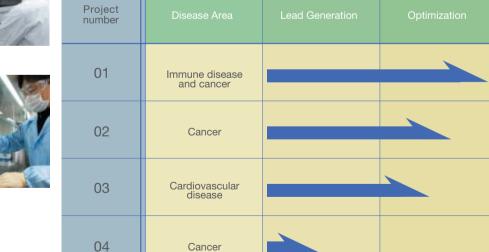
## Kinase Inhibitor Programs



Carna Biosciences is building an internal drug pipeline through the application of our key technologies. We have successfully applied our proprietary kinase selectivity profiling, QuickScout® to generate novel, potent and selective lead compounds within a few months for many kinase targets. We are taking a rational approach to optimize a lead compound into drug candidates using X-ray co-crystallography information and computer-aided drug design. Our strategy for the further development of kinase inhibitors is to profile compounds against a panel of 276 kinases, and we believe this approach lead to a better understanding of structure-activity relationships, helping to avoid targets with toxicity and to select better drug candidates.



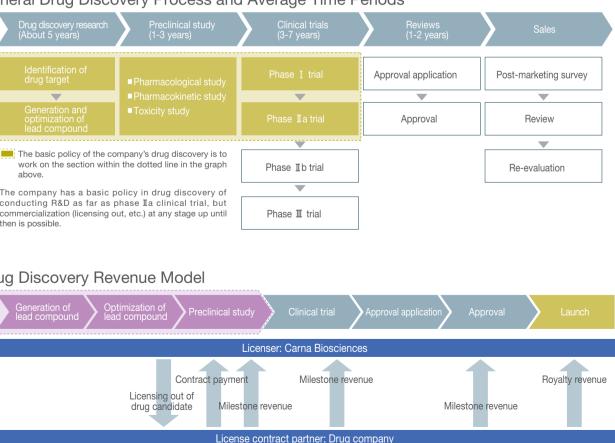
#### **Drug Discovery Project** Research Pipelines for Kinase inhibitors



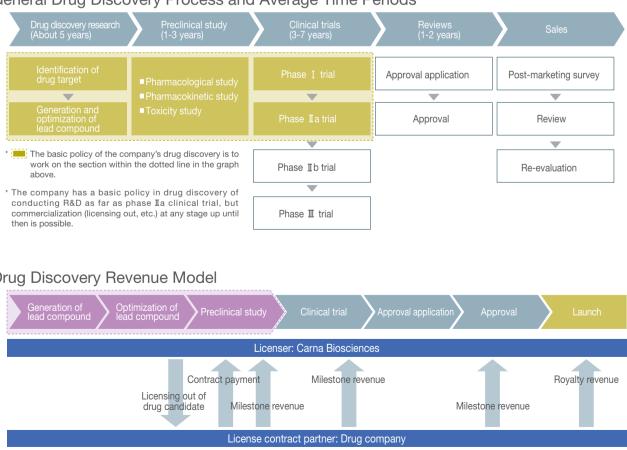
### Revenue Model

Carna Biosciences has a basic policy in drug discovery of conducting R&D as far as phase IIa clinical trial and considers outsourcing drug candidates to pharmaceutical companies at any of the stages prior to that. The company has envisioned a revenue model whereby outsourcing new drug candidates to pharmaceutical companies allows it to receive contract payments upon the conclusion of licensing contracts, milestone revenue based on licensing contracts for payment prior to entering each clinical trial, upon approval application and upon approval acquisition, and fixed rate royalties for sales or net sales of new drugs following their launches.

### General Drug Discovery Process and Average Time Periods



#### Drug Discovery Revenue Model



\* 🛄 The basic policy of the company's drug discovery is to work on the section within the dotted line in the graph above.







#### Balance Sheets

Balance Sheets Thousands o		
	<b>FY2006</b> As of Dec. 31, 2006	<b>FY2007</b> As of Dec. 31, 2007
Assets		
Current assets Fixed assets Tangible fixed assets Intangible fixed assets Investments and other assets Total assets Liabilities	1,589,488 349,088 91,670 1,826 255,592 1,938,577	1,351,810 270,565 84,117 7,458 178,989 1,622,375
Current liabilities	185,982	148,609
Long-term liabilities Total liabilities	99,706 285,689	37,846 186,455
Net Assets		
Shareholders' equity Common stock Capital surplus Retained earnings Valuation and translation adjustments Total net assets Total liabilities and net assets	1,595,267 1,521,700 1,467,500 -1,393,932 57,620 1,652,888 1,938,577	1,415,438 1,521,700 73,567 -179,829 20,481 1,435,920 1,622,375

#### Statements of Income

Thousands of Income Thousands of Income		
	<b>FY2006</b> Jan. 1, 2006 - Dec. 31, 2006	<b>FY2007</b> Jan.1 , 2007 - Dec. 31, 2007
Net sales	405,002	513,922
Cost of sales	98,705	99,136
Gross profit	306,296	414,786
Selling, general and administrative expenses	562,597	562,204
Operating loss	256,300	147,418
Non-operating income	5,843	8,698
Non-operating expenses	19,314	20,142
Ordinary loss	269,771	158,861
Extraordinary income	131	—
Extraordinary loss	62,365	19,963
Loss before income taxes	332,006	178,825
Income taxes	950	1,004
Net loss	332,956	179,829

### Statements of Cash Flows

	<b>FY2006</b> Jan. 1, 2006 - Dec. 31, 2006	<b>FY2007</b> Jan. 1, 2007 - Dec. 31, 2007
Net cash provided by (used in) operating activities	-181,498	-192,603
Net cash provided by (used in) investing activities	-86,383	-66,228
Net cash provided by (used in) financing activities	1,521,701	-50,000
Effect of exchange rate changes on cash and cash equivalents	80	846
Increase in cash and cash equivalents	1,253,899	-307,984
Cash and cash equivalents at beginning of year	255,115	1,509,014
Cash and cash equivalents at end of year	1,509,014	1,201,029



Profile	
Name:	Carna Biosciences, Inc.
Address:	KIBC 511, 5-5-2
	Minatojima-Minamimachi, Chuo-ku,
	Kobe 650-0047, Japan
Phone:	+81-78-302-7039
Fax:	+81-78-302-6665
Established:	April 10, 2003
Capital:	1,964,570,000 yen (as of August 31, 2008)
Number of Employees:	42 (as of June 30, 2008)

Overseas Subsidia	ary
Name:	CarnaBio USA, Inc.
President:	Takashi Hara
Address:	209 West Center Street, Suite 127, Natick, MA 01760, USA

### History

Thousands of yen

JUN 2008	Concluded a joint research agreement w
APR 2008	Established US subsidiary CarnaBio US
MAR 2008	Listed on JASDAQ NEO securities excha
OCT 2007	Concluded a collaboration and services services with OSI Pharmaceuticals, Inc.
OCT 2007	Newly established a chemistry laborate (HI-DEC) aiming at further acceleration in
APR 2007	Concluded a joint research agreement joint research on new drug candidate co
JUL 2006	Concluded a distributorship agreement
FEB 2006	Entered into collaboration with SBI Biot joint research on new drug discovery on
DEC 2005	Concluded an exclusive Japanese dist kinase proteins
AUG 2004	Newly established research office at BL (BMA, Kobe) and started animal experime
JUN 2004	Entered into collaboration with Crystal discovery on target kinases
APR 2003	Spin-off from Nippon Organon K.K. to e

Board Members and Auditors		
President and CEO:	Kohichiro Yoshino	
Chairman:	Norihiro Tsugi	
Director:	Hiroshi Ishiguro	
Director:	Norio Aikawa	
Director:	Yutaka Shimakawa	
Director:	Takashi Hara	
Director:	Satoru lino	
Auditor (standing):	Astuo Arita	
Auditor:	Tsuguo Ogasawara	
Auditor:	Kiyoshi Nakai	

Phone:	+1-508-650-1244
Fax:	+1-508-650-1722
Established:	June 9, 2008
Capital:	400,000 U.S. dollars

with National Cancer Center for anti-cancer drug

SA, Inc. in Boston suburb

hange

as agreement for a joint venture in lead compound discovery (USA)

tory at the Kobe Healthcare Industry Development Center in drug discovery research

with SBI Biotech Co., Ltd. and Crystal Genomics Inc. for compound development (2nd step)

with KinaseDetect ApS of Denmark

otech Co., Ltd. and Crystal Genomics Inc. (South Korea) in n target kinases (1st step)

stributorship agreement with Toyobo Co., Ltd. for sales of

usiness Support Center for Biomedical Research Activities ents for initial evaluation of low molecular compounds

Genomics Inc. (South Korea) in joint research on new drug

establish Carna Biosciences, Inc. on Port Island, Kobe