## Financial Results for the 2nd Quarter of Fiscal Year 2022

(April 1, 2022 – September 30, 2022)



## SEIKAGAKU CORPORATION

(TSE: 4548)

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#### < Cautionary Notes >

This material contains forward-looking statements regarding future management strategies or performance forecasts. These statements are based on judgments derived from information that is currently available to Seikagaku and are subject to risk and uncertainty. Actual results and developments may differ significantly from these forward-looking statements due to various factors.

Information about pharmaceutical products or medical devices (including products currently in development) included in this material is not intended to constitute an advertisement or medical advice.



## **Overview for 2Q of FY 2022**

	2Q FY2022 Results			Yea	ar-o	n-Year	
(Millions of Yen)				Change		% of Change	
Net sales		17,258		-3,2	272		-15.9%
Operating Income		2,610		-3,4	28		-56.8%
Ordinary Income		3,600		-2,7	<b>'93</b>		-43.7%
Net Income		3,141		-1,8	877		-37.4%
<b>R&amp;D Expenses</b> (excluding royalty income)		<b>3,407</b> (19.7%)		•	<b>94</b> 7pt)		<b>-10.4</b> %
Average Exchange Rate (1US\$)		¥ 133.98		+24	4.18		
		2Q FY2022 Results		2Q FY2021 Results		Reference) 22 Forecasts	
Net Income per Share		¥56.3	32	¥89.07		¥48.12	



## Net sales by Business Segment (2Q of FY2022)

(N	Aillions of Yen)	2Q FY2022 Results	Year-on- Year	% of Change
	Net sales	17,258	-3,272	-15.9%
Р	harmaceuticals	11,692	-4,444	-27.5%
	Domestic Pharmaceuticals	5,808	-997	-14.7%
	Overseas Pharmaceuticals	4,384	-65	-1.5%
	Bulk Products /CDMO	1,497	+167	+12.6%
	Royalty Income	1	-3,548	-100.0%
L	AL Business	5,566	+1,171	+26.7%
(0)	verseas sales)	10,353	+1,218	+13.3%

\* Foreign exchange impact on overall net sales : approx. +1,250million yen

## **Domestic Pharmaceuticals**

#### ► ARTZ (Joint-function improving agent)

• Shipment volume up due to shipment timing, but sales down due to the impact of NHI drug price reductions

#### **JOYCLU** (Joint-function improving agent)

• Substantial decline in sales reflecting concentration of shipments in 1H FY2021 because of the launch

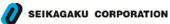
#### • **OPEGAN series** (Ophthalmic viscoelastic devices)

• Sales down due to a sales partner inventory adjustment and NHI drug price reductions

#### MucoUp

#### (Submucosal injection agent for endoscopic surgery)

- Sales down, reflecting a decline from high level of shipments in first half of previous fiscal year and impact of NHI reimbursement price revision
- HERNICORE (Treatment for lumbar disc herniation)
  - Sales up as a result of the shipment schedule



## Net sales by Business Segment (2Q of FY2022)

(Millions of Yen)	2Q FY2022 Results	Year-on- Year	% of Change
Net sales	17,258	-3,272	-15.9%
Pharmaceuticals	11,692	-4,444	-27.5%
Domestic Pharmaceuticals	5,808	-997	-14.7%
Overseas Pharmaceuticals	4,384	-65	-1.5% -
Bulk Products /CDMO	1,497	+167	+12.6%
Royalty Income	1	-3,548	-100.0%
LAL Business	5,566	+1,171	+26.7%
(Overseas sales)	10,353	+1,218	+13.3%

\* Foreign exchange impact on overall net sales : approx. +1,250million yen

#### **Overseas Pharmaceuticals**

\*Foreign exchange impact on Overseas Pharmaceuticals: approx. +500 million yen

- ► Gel-One in the U.S. (Single injection)
  - Sales substantially up, fueled by the impact of yen appreciation



Gel-One: Single-injection joint function improving agent

## **SUPARTZ FX in the U.S.** (Multiple injection)

• Sales down due to shipment timing

#### ARTZ in China (Multiple injection)

- Sales down substantially, with no shipments through July due to a packaging material change
- Resumption of shipments in August, on schedule



## Net sales by Business Segment (2Q of FY2022)

(Mi	illions of Yen)	2Q FY2022 Results	Year-on- Year	% of Change	Bulk Products / CDMO Bulk products sales at the prior-year level, and CDMO
٦	Net sales	17,258	-3,272	-15.9%	sales at Dalton Pharmaceutical up due to the impact of yen depreciation *Foreign exchange impact on Bulk Products/CDMO:
Ph	armaceuticals	11,692	-4,444	-27.5%	Royalty Income approx. +120million yen
	Domestic Pharmaceuticals	5,808	-997	-14.7%	Substantial decline LAL Business
	Overseas Pharmaceuticals	4,384	-65	-1.5%	In addition to a steady trend in domestic sales, sales up on growth in sales of ACC's reagents for Bacterial Endotoxin Testing (BET), contract services and
	Bulk Products CDMO	1,497	+167	+12.6%	Clinical Diagnostic (Fungitell) reagents *Foreign exchange impact on LAL Business: approx. +630million yen
F	Royalty Income	1	-3,548	-100.0%	Overseas Sales Ratio(excluding royalty income)
LA	L Business	5,566	+1,171	+26.7%	Domestic 40.0% Year-on-Year
(Overseas sales)		10,353	+1,218	+13.3%	Overseas +6.2pt

\* Foreign exchange impact on overall net sales : approx. +1,250million yen

■ Overseas LA, Bulk/CDMO ■ Overseas Pharmaceuticals ■ Domestic sales



## Income for 2Q of FY2022

		2Q FY2022		% of	<b>Operating Income</b>	2,610 (-3,428)			
(Millions of Yen)		Results	Year	Change	Cost of sales ratio (-0.3pt)				
	Net sales	17,258	-3,272	-15.9%	<ul> <li>Ratio at the prior-year level because of change sales mix, despite NHI drug price reductions</li> <li>SGA Expenses (+86)</li> <li>R&amp;D expenses down due to completion of</li> </ul>				
	Cost of Sales (excluding royalty income)	<b>7,427</b> (43.0%)	+ <b>69</b> (-0.3pt)	+0.9%					
	SGA expenses	7,220	+86	+1.2%	in the U.S. for SI-6603				
	R&D Expenses (excluding royalty income)	<b>3,407</b> (19.7%)	<b>-394</b> (-2.7pt)	-10.4%	<ul> <li>Higher expenses at or including foreign curr</li> <li>Ordinary Income</li> </ul>				
	perating Income to Net sales ratio)	<b>2,610</b> (15.1%)	-3,428 (-14.3pt)	- <b>56.8</b> %	• Increase in foreign ex	ne / Expenses (+635)			
	Ordinary Income	3,600	-2,793	-43.7%	Net Income	<b>3,141(-1,877)</b>			
Net Income		3,141	-1,877	-37.4%	Income Taxes (-915)				
	Depreciation	610	+135	+28.6%	<ul> <li>Decline from high profits in previous fisca</li> </ul>				



## **Overview of Forecasts in FY2022**

	FY2	2022	FY2021 Results							Degree of	
(Millions of Yen)	Forecasts		R	Results Cha		hange	nge % of Change		nge	Progress	
Net sales	3	3,500		34,851		-1,3	351	-3	.9%	51.5%	
Operating Income		1,700		4,495		-2,7	795	-62	.2%	153.6%	
Ordinary Income		2,900		5,395		-2,4	495	-46	.2%	124.2%	
Net Income		2,650		3,733		-1,(	083	-29	.0%	118.6%	
R&D Expenses (Ratio to net sales excluding royalty income)		<b>8,000</b> (23.9%)		<b>9,005</b> (29.2%)		- <b>1,(</b> (-5.	<b>)05</b> .3pt)	-11	.2%	42.6%	
Average Exchange Rate (1US\$)	2Q forecast	¥140.00		¥112.38							
		FY202 Forecas			Exchange			Rate Sensitivity ge of ¥1against the US\$)			
Net Income per	share	¥4	8.12	¥6	6.32		Net sales Ap		Арр	orox. ¥45 million	
Dividend per share Dividend Payout ratio			26.00 4.0%			0	Operating income App		Арр	prox. ¥15 million	



## **Forecasts (Net sales) in FY2022**

FY2022		Year-on-Year		Net sales		
(Millions of Yen)	Forecasts	Change	% of Change	Forecasting a sales decrease to result from steep		
Net sales	33,500	-1 351 -3.9% price reductions, de		decline in royalty income and impact of NHI drug price reductions, despite higher pharmaceutical sales volumes in Japan and positive impact of yen		
Pharmaceuticals	22,500	-3,196	<b>-12.4</b> %			
Domestic Pharmaceuticals	10,850	-597	-5.2%	<b>Domestic pharmaceuticals :</b> Sales decrease to result from NHI price reductions		
Overseas Pharmaceuticals	8,650	+997	+13.0%	<b>Overseas pharmaceuticals :</b> Sales increase due to yen depreciation		
Bulk Products /CDMO	3,000	+392	+15.1%	Bulk products/CDMO : CDMO sales increase due to yen depreciation		
Royalty income	1	-3,988	-100.0%	Royalty income : Substantial decline		
LAL Business	11,000	+1,844	+20.2%			
(Overseas sales)	20,650	+2,731		<b>LAL Business</b> Expected sales increase due to the impact of yen depreciation on overseas sales		
* Forei	gn exchange imp	act on overall approx. +3,100	net sales : )million yen			

SEIKAGAKU CORPORATION

## Forecasts (Income) in FY2022

	FY2022	Year-on	-Year
(Millions of Yen)	Forecasts	Change	% of Change
Net sales	33,500	-1,351	-3.9%
OperatingIncome (Ratio to net sales)	<b>1,700</b> (5.1%)	-2,795 (-7.8pt)	-62.2% -
Ordinary Income	2,900	-2,495	-46.2%
Net Income	2,650	-1,083	-29.0%
<b>Cost of Sales ratio</b> (excluding royalty income)	47.2%	+ <b>0.8</b> pt	-
R&D Expenses	8,000	-1,005	-11.2%
R&D Expenses ratio (excluding royalty income)	23.9%	- <b>5.3</b> pt	-
Depreciation	1,300	+248	+23.6%

#### **Operating Income**

#### **SGA** expenses

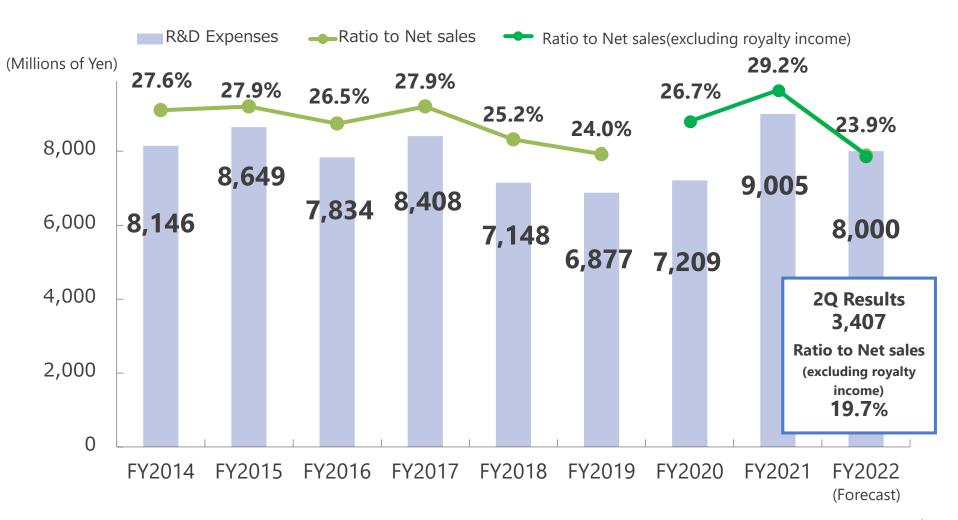
- Decrease in R&D expenses accompanying completion of subject enrollment in an additional clinical study in the U.S. for SI-6603
- Higher expenses at overseas subsidiaries, including foreign currency translation

#### **Ordinary Income**

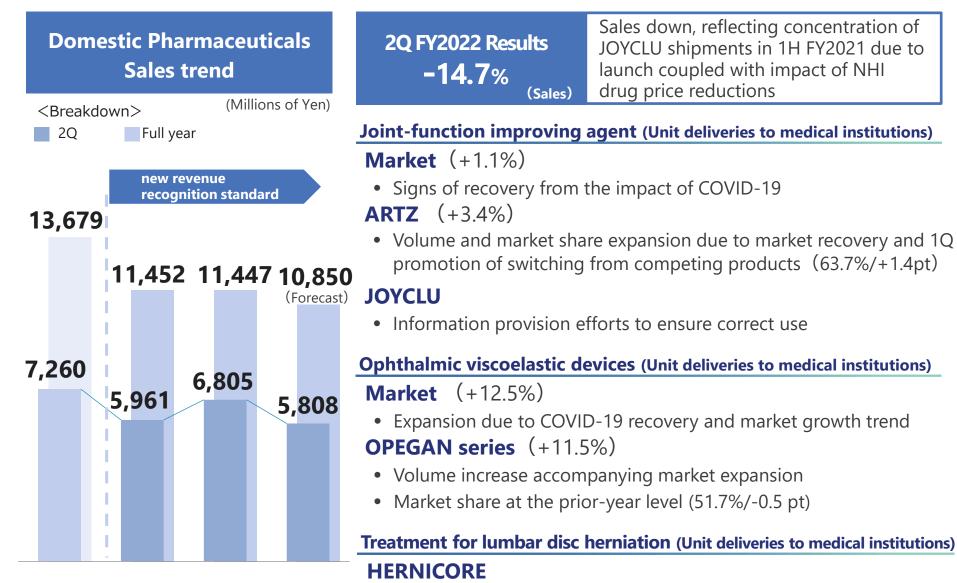
#### **Non-operating Income/Expenses**

 Increase in foreign exchange gains related to valuation of foreign currency-denominated assets

## Continuing to focus on drug discovery specializing in Glycoscience Undertaking improvement in R&D efficiency



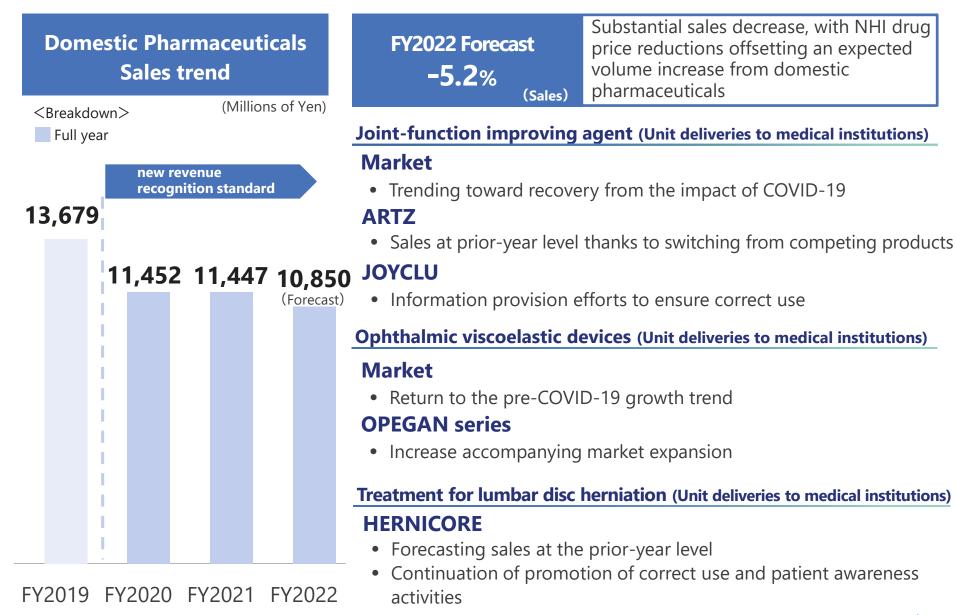




- FY2019 FY2020 FY2021 FY2022 .
  - Sales at the prior-year level

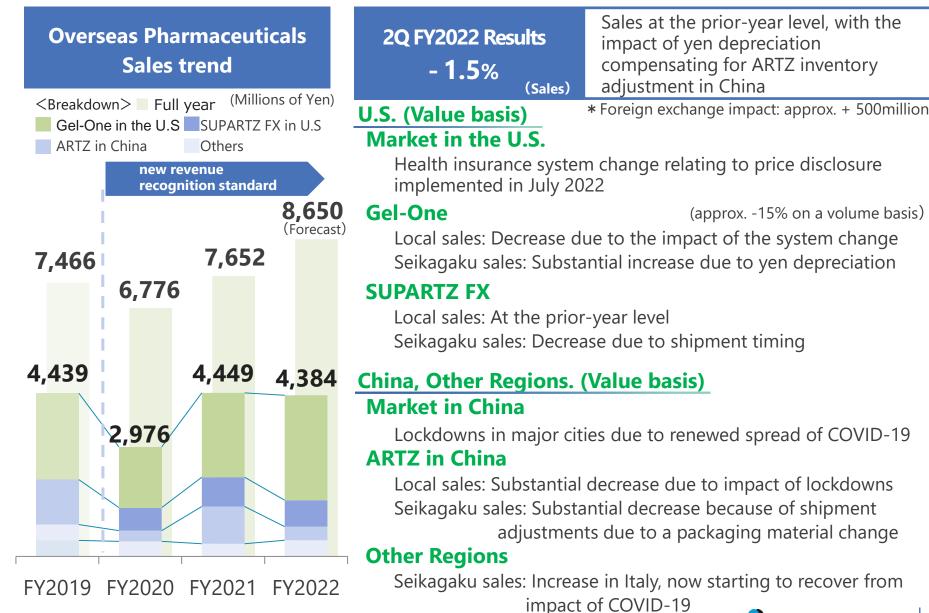


## Domestic Pharmaceuticals (FY2022 Forecast Year-on-Year)





## **Overseas Pharmaceuticals (Year-on-Year)**



## **Overseas Pharmaceuticals (FY2022 Forecast Year-on-Year )**

Sales increase, with impact of yen **Overseas Pharmaceuticals** FY2022 Forecast depreciation compensating for impact of system changes in the U.S. and ARTZ Sales trend +13.0% shipment adjustments in China (Sales) (Millions of Yen) <Breakdown> Full year \* Foreign exchange impact: approx. + 1,200million U.S. (Value basis) Market in the U.S. new revenue Expected impact from system change relating to price disclosure recognition standard **Gel-One** 8,650 Local sales: Decrease due to impact of system change (Forecast) Seikagaku sales: Increase because of impact of yen depreciation 7,652 7,466 **SUPARTZ FX** 6,776 Local sales: Decrease due to trend toward preference for products requiring few injections Seikagaku sales: Increase because of impact of yen depreciation China, Other Regions. (Value basis) Market in China Expected impact from lockdowns due to new outbreaks of COVID-19 **ARTZ in China** Local sales: Substantial decline due to lockdowns Seikagaku sales: Decrease because of shipment adjustments due to a packaging material change **Other Regions** 

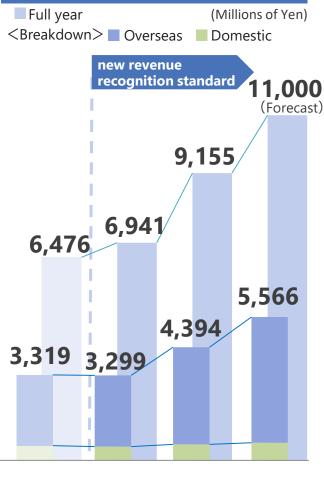
FY2019 FY2020 FY2021 FY2022

Seikagaku sales: Increase in Italy, now starting to recover from impact of COVID-19



## Sales of LAL Business (year-on-year / value basis)

#### LAL Business Sales trend



FY2019 FY2020 FY2021 FY2022

#### 2Q FY2022 Results : +26.7% (Year-on-Year)

\* Foreign exchange impact: approx. + 630million

#### **Overseas**

Sales increase on growth from Bacterial Endotoxin Testing (BET) reagents, Clinical Diagnostic (Fungitell) reagents, and contract testing services at overseas subsidiary ACC

## Domestic

Sales increase on solid sales of reagents and other products

#### FY2022 Forecasts : +20.2% (Year-on-Year)

\* Foreign exchange impact: approx. + 1,650million

### **Overseas**

Sales increase due to the impact of yen depreciation

## Domestic

Forecast of steady sales

AL Business :	The manufacturing and sales of Endotoxin detection reagents* used in the quality control of pharmaceuticals and medical devices

\*Endotoxin detection reagents are reagents whose main ingredient is Limulus Amebocyte Lysate (LAL).



## Mid-term Management Plan FY2022 to FY2025





- Summary of Previous Mid-term Management
- Overview of Current Mid-term Management Plan

## Key Measures

- I . Accelerate R&D utilizing unique drug-discovery technologies
- II. Maximize the product value of SI-6603 (treatment for lumbar disc herniation)
- II. Maintain and enhance the business value of joint function improving agents
- IV. Construct a global production system
- V. Expand the LAL business through recombinant technologies

## • Sustainability

## • Numerical Targets and Basic Policy on Profit Distribution

- Numerical Targets
- Basic Policy on Profit Distribution



#### I. Accelerating new drug discovery to become the pillar of new profits

- Joint function improving agent JOYCLU launched
- SI-722 (treatment for interstitial cystitis) and SI-449 (adhesion barrier) advanced to the next stage of development
- Enrollment for SI-6603 (treatment for lumbar disc herniation) completed
- SEIKAGAKU NORTH AMERICA CORPORATION established

## **I**. Solidifying the profit foundation through market expansion of new products

- HyLink (intra-articular single-injection viscosupplement for the treatment of knee osteoarthritis) launched in Taiwan
- Alliance agreement concerning SI-613 concluded with Eisai (China, South Korea)
- PyroSmart NextGen Recombinant LAL Reagent launched

#### **III. Productivity improvement reforms**

- DALTON CHEMICAL LABORATORIES, INC. made a subsidiary
- Business continuity system developed in response to the impact of COVID-19
- Progress with review of procurement costs and sales-related expenses

We achieved a measure of success with each of the three important measures in the management plan, successfully solidifying the profit foundation to return Seikagaku to a growth trajectory



## **Previous Mid-term Management Plan Numerical Targets**

	FY2021 results*	FY2021 target (disclosed Nov. 2019)	Percentage achievement
Net sales	¥31.2 billion	¥28.3 billion	+10.5%
Ordinary income	¥5.39 billion	¥4.5 billion	+19.9%
SKK EBITDA	¥5.54 billion	¥5.0 billion	+10.9%
<b>Overseas sales ratio</b> (excluding royalty income)	56.6%	50.0%	+6.6 pt.

\*Converted to previous presentation categories SKK EBITDA: A profit indicator that adds depreciation and royalty income to operating income



## We also achieved each numerical target in the plan







#### **Business Environment**

Pharmaceutical Industry Trends		Societal Trends	
Domestic pharmaceutical market		Development of international frameworks and principles	
• Growth in healthy longevity	<ul> <li>Diversification of treatment options</li> <li>Impact of drug price system</li> <li>Strengthening of quality assurance and control systems</li> </ul>	<ul> <li>Adoption of the SDGs by the U.N., global penetration</li> <li>Adoption of the Paris Agreement at COP21</li> </ul>	
Overseas joint function improving agent market	R&D environment	Revision of the Corporate     Governance Code	
<ul> <li>Osteoarthritis patients increasing (aging, etc.)</li> <li>Greater numbers of active seniors</li> <li>Intensification of competition due to new product introductions, etc.</li> <li>A broad range of treatment options</li> <li>System change relating to price disclosure</li> </ul>	<ul> <li>Increasing R&amp;D costs and risks</li> <li>Depletion of seeds for drug discovery</li> <li>Appearance of new treatment methods, such as regenerative medicine</li> <li>Diversification of new drug modalities</li> </ul>	Stakeholder expectations	
		<ul> <li>Increase in sustainable investment ratios</li> <li>Growing importance of non-financial information disclosure in investment decisions</li> </ul>	

A flexible response to an era of extreme environmental change and fulfillment of social responsibility are required





## **Overview of Current Mid-term Management Plan**

Exploring the Innovative Promise of Glycoscience



Based on a solidified profit foundation, nurture the capability to maintain a constant growth trajectory by energetically implementing the key measures <u>Aim to achieve record-high business results in the final year of the plan</u>

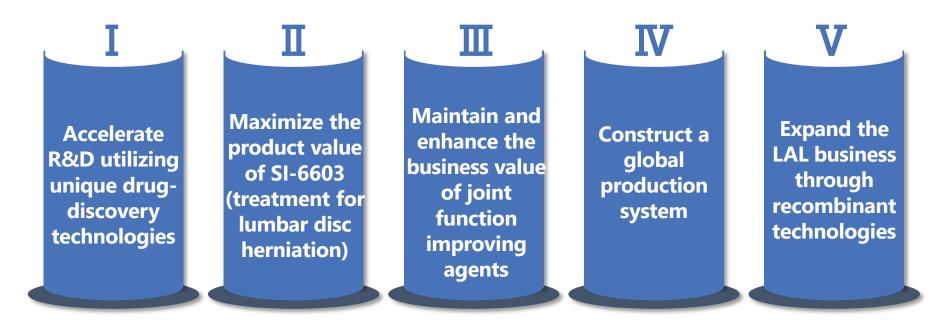












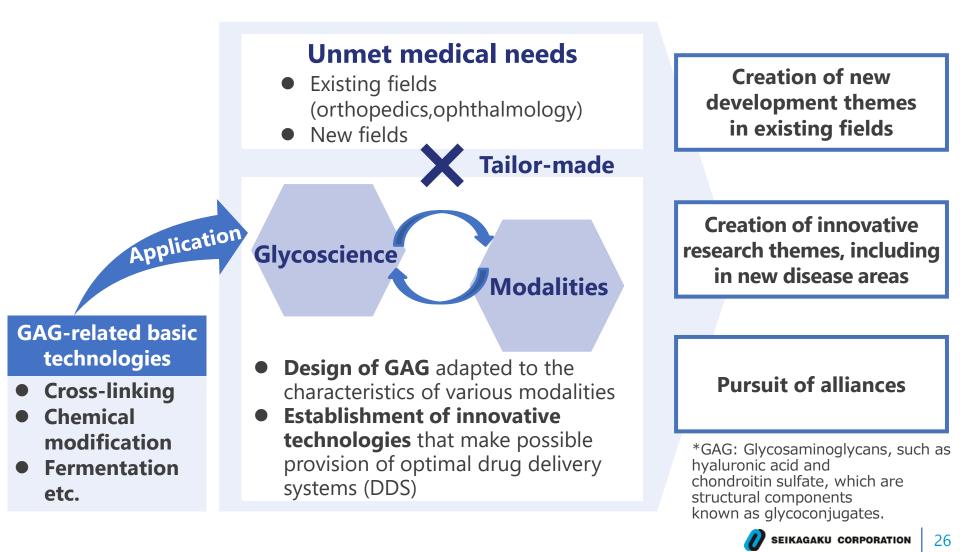
#### **Business foundation**

Enhancement of employee engagement/ Organizational strengthening and human resource development





# Aim to leverage Seikagaku's GAG-related technologies to create products needed by patients



#### I. Accelerate R&D Utilizing Unique Drug-Discovery Technologies

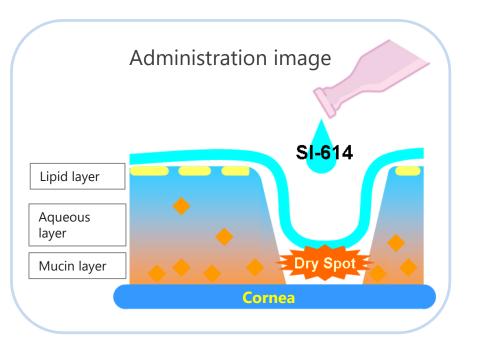
## **Pipeline List (Research and Development themes)**

**Exploring the Innovative** Promise of Glycoscience

Development code/ Indication (Product name)	PI PI PII Appli- Market cation approval	
<b>SI-6603</b> Lumbar Disk Herniation (Condoliase)	SI-6603 US US US	
<b>SI-614</b> Dry eye (Modified Hyaluronate)	SI-614 US	
<b>SI-613</b> US : Knee Osteoarthritis (Diclofenac etalhyaluronate sodium)	SI-613 US	
<b>SI-613-ETP</b> Enthesopathy (Diclofenac etalhyaluronate sodium)	SI-613- ETP JP PIb ※ Discontinued (February 2022)	
<b>SI-722</b> Interstitial cystitis and bladder pain syndrome (Steroid compound sodium chondroitin sulfate)	SI-722 US PI/I	
Development code/ Indication (Product name)	Pilot Pivotal Application Market approval	
<b>SI-449</b> Adhesion Barrier (Cross-linked Chondroitin Sulfate)	SI-449 JP JP	
SI-449 US : Adhesion Barrier (Cross-linked Chondroitin Sulfate)	SI-449 US	
(Cross-linked Chondroitin Sulfate) SI-449 US : Adhesion Barrier	SI-449	



#### Phase III clinical study in the U.S. initiated in May 2022 Aiming to complete the Phase III study during the period of the mid-term management plan



#### **Development status**

- ► U.S. : P III
  - **Initiated May 2022**
  - Clinically beneficial effects confirmed in PII/PIII clinical studies
  - Evaluation of efficacy and safety

## **Expected Features**

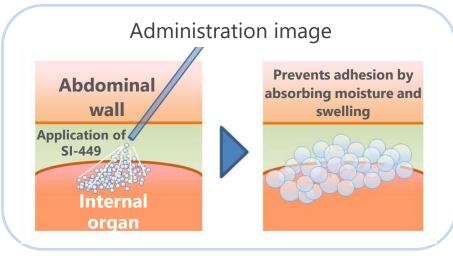
- SI-614 Improves symptoms of dry eye by protecting the ocular surface and promoting corneal epithelial wound healing
- Dry Eye is a multifactorial disease, and Seikagaku aims to provide a treatment option based on a new mechanism unavailable from existing products

<SI-614 summary>
Dev. Code : SI-614 Generic name : Modified Hyaluronate
Product name : Dry eye Formulation : Ophthalmic solution
Estimated U.S. patients : 14 million (Seikagaku estimates)



## SI-449 (Adhesion Barrier / Medical Device)

## Subject enrollment for pivotal study completed in September 2022 Aiming to obtain approval and initiate a clinical study in the U.S. during the mid-term management plan period



## **Expected Features**

Leveraging pulverized characteristics to respond to healthcare needs through the spreading use of laproscopic surgery

## **Development status**

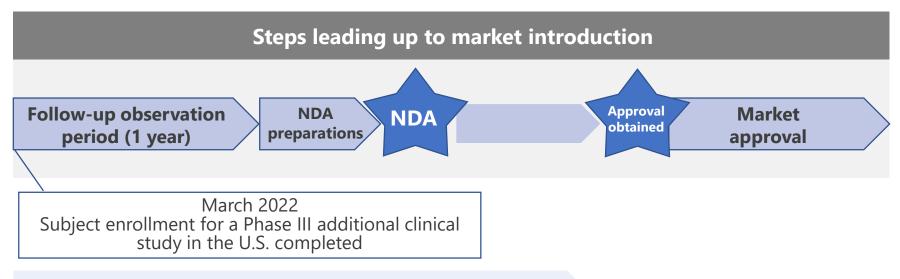
- Japan pivotal study(field of gastroenterological surgery) Starting May 2020
  - Evaluated for effectiveness, safety, and usability
     ⇒Subject enrollment complete in Sept. 2022
- Japan pilot study (field of gynecology) Starting November 2021
  - Confirming operability and safety
  - Aiming to expand range of applications
  - ⇒Subject enrollment complete in May 2022
- Proceed with development with a view to global development; Start of U.S. study under review

<SI-449 summary> Dev. Code : SI-449 Generic name : Cross-linked chondroitin sulfate Product name : Adhesion barrier Method of use : Intra-abdominal application (powdered formulation) Adhesion barrier market : Japan: ¥14 billion, Global: ¥100 billion (Seikagaku estimates)





#### Obtain clinical study findings during FY2025 Aim for an NDA and approval during the period of the current mid-term management plan

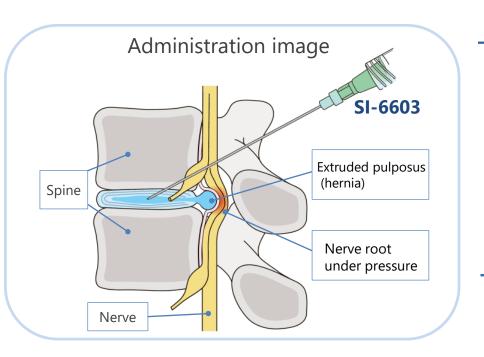


- Cooperation with SNA\* in prompt and accurate NDA and response to regulatory review to obtain approval
- Timely, close communication with the regulatory authorities
  - Sales preparations in cooperation with the sales partner
  - Construction of a stable supply system

\* SEIKAGAKU NORTH AMERICA: Established in January 2022 in Canada with the aim of accelerating development of pharmaceuticals and medical devices and obtaining regulatory approval



## Aim to provide new options in lumbar disc hernia treatment and maximize product value through early market penetration



## **Expected Features**

- Relieving symptoms by decreasing intradiscal pressure and reducing the pressure on the nerve root by single injection
- It can be administered without general anesthesia, and the administration can be less invasive for the patient compared to surgical technique because of direct intradiscal injection

## Estimated U.S. patients

New patients with lumbar disc herniation : 3 to 5 million per year (Seikagaku estimates)

<SI-6603 summary> Dev. Code : SI-6603 Generic name : Condoliase Indication : Lumbar disc herniation Method of use : Injection into Iumbar disc (under X-ray observation)



III. Maintain and Enhance the Business Value of Joint Function Improving Agents Maintain and Enhance the Business Value of Joint Function Improving Agents

Exploring the Innovative Promise of Glycoscience

Pursue corporate value enhancement through maximization of Seikagaku products in the market for joint function improving agents and cost structure improvement

Maximization of Seikagaku products in the market for joint function improving agents

ARTZ

 Share expansion through switching from competing products JOYCLU

- Conduct of clinical research to identify the cause of shock and anaphylaxis
- Continuation of information provision to ensure appropriate treatment and measures

#### **Cost structure improvement**

- Packaging material specification change
- Manufacturing process efficiency improvement

# Maintenance and enhancement of business value that supports business management





## Safety Information on Joint Function Improving Agent JOYCLU

Blue Letter (rapid safety information) issued in June 2021 Seikagaku is further raising awareness among healthcare professionals and taking measures to identify the cause

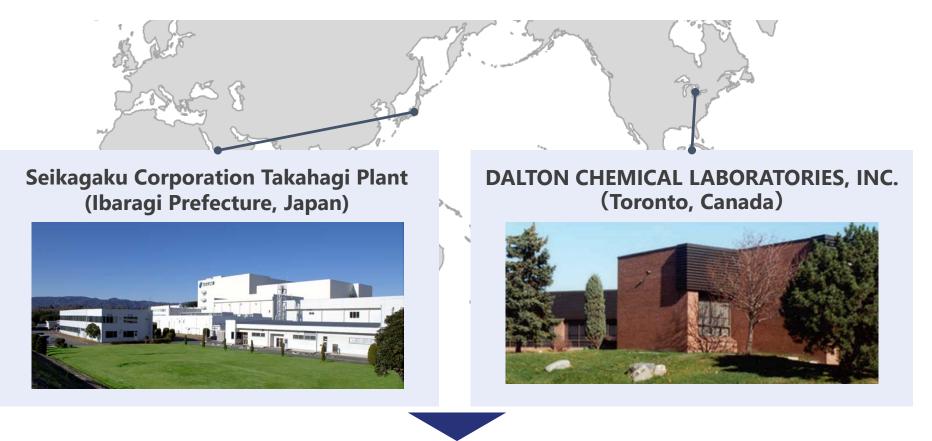
**Dear Healthcare Professionals Letter of Rapid Safety Communication (Blue Letter)** issued on June 1 in response to multiple reports of anaphylaxis occurring in patients following administration of JOYCLU

- To ensure patient safety by promptly alerting healthcare professionals of the situation to enable the provision of appropriate treatment and measures
- Addition of a WARNING section to the drug package insert and revision of the IMPORTANT PRECAUTIONS section
- Leaflet created to alert patients and their families to the risk of side effects
- Continued gathering of safety information and provision of safety-related information
- Physician-led clinical research to identify the cause initiated in April 2022

Continue gathering of safety information and provision of safety-related information in collaboration with sales partner Ono Pharmaceutical Aim to contribute to appropriate treatment on the basis of clinical research findings



## Further reinforce a stable supply of products on the basis of a GMP-compliant, appropriate governance system



Undertake construction of a global production system with Japan and North America as dual production bases, including transfer of production of some products



# Aim to be a leading company in the market for recombinant endotoxin detection reagents and glucan detection in vitro diagnostics

#### **Pursuit of Unique Value**

Refine the joint development system on the basis of further collaboration with ACC.\*

Next-generation products (recombinant technologies, new reagents, new measurement systems)

#### **Collaboration with Affiliated Companies**

Create value from business models and operating systems that cannot be created independently

Automatic measurement devices



\* Associates of Cape Cod: Overseas subsidiary responsible for overseas expansion of the LAL business

Creation of unique value that competitors cannot imitate in a market with strong customer needs



## Accomplish a transformation necessary from a long-term perspective in addition to continuation of current businesses and product improvement

## **1. Endotoxin detection reagent business**

- Acceleration of global expansion of endotoxin detection reagents leveraging recombinant technologies
- Accumulation of reliable scientific data utilizing PyroSmart NextGen Recombinant LAL Reagent (launched in 2021) and market penetration and sales promotion through support for early adopters

### 2. Diagnostics business

- Increase in the number of countries where glucan detection in vitro diagnostics are sold and acceleration of new expansion into the hospital market
- Progress in development of recombinant glucan detection in vitro diagnostics and product portfolio expansion

### 3. Measurement equipment and software business

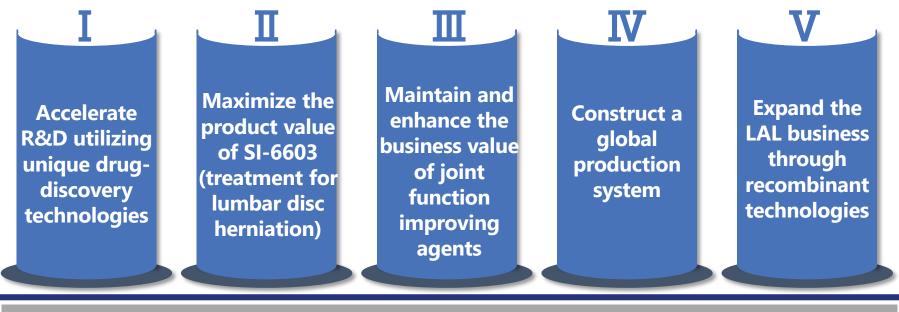
 Promotion of development and improvement of automatic measurement equipment and software in Japan and overseas in collaboration with an affiliated company

## Key Measures Overview of Key Measures -Business Foundation

Exploring the Innovative Promise of Glycoscience

Introduction of a new HR system in October 2022 Work to solidify the business foundation for achieving sustained growth through enhancement of employee engagement as well as organizational strengthening and human resource development

## **Five Key Measures (Strategic Pillars)**



### **Business foundation**

Enhancement of employee engagement/ Organizational strengthening and human resource development



**Exploring the Innovative Promise of Glycoscience** 







Seikagaku has identified important issues that should be addressed on a priority basis in the interest of achieving sustainable development of society and enhancement of corporate value





Seikagaku will focus on initiatives related to these material issues, which will become the foundation for the key measures in the mid-term management plan



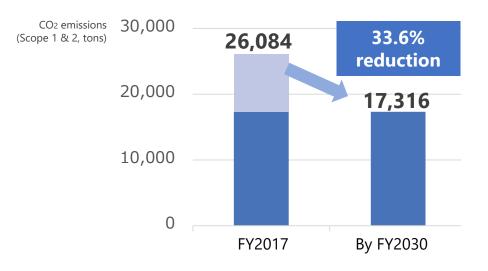
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Healthcare	<ul> <li>Acceleration of multinational expansion and new drug development.</li> <li>Strengthening and utilization of original GAG-related basic technologies</li> <li>Product improvement and post-marketing product development aligned with medical needs</li> <li>Related material issues : ①、②、③</li> </ul>
Environment	<ul> <li>CO2 emissions reduction (33.6% reduction from the reference year by FY2030)</li> <li>Zero waste emissions and promotion of use of recycled raw materials</li> <li>Biodiversity preservation activities, including reduction or elimination of use of animal-derived materials in products</li> <li>Related material issues : 6</li> </ul>
Society	<ul> <li>Strengthening of initiatives for respect of human rights in the supply chain</li> <li>Boosting of employee motivation through strengthened HR systems and introduction of diversity measures and work-life balance measures that lead to employee retention</li> <li>Related material issues : 5</li> </ul>
Governance	<ul> <li>Development and operation of a highly effective corporate governance system</li> <li>Fostering of compliance awareness and a quality culture</li> <li>Enhancement of preventive maintenance activities for a stable supply of products</li> <li>Related material issues : ④</li> </ul>



## Initiate measures during the period of the current mid-term management plan aimed at reducing CO<sub>2</sub> by 33.6% from the FY2017 level by FY2030



# 1. Energy saving

- Conversion to LED lighting at plants and laboratories
- Review of air conditioner operation, such as introduction of nighttime mode

# 3. Updating of equipment

 Modification to the filtration method used for sterile water for injection production equipment

# 2. Fuel conversion

Conversion of boiler fuel to liquid natural gas (LNG)

## 4. Introduction of renewable energy

• Introduction of solar power and other renewable energy sources



**Exploring the Innovative Promise of Glycoscience** 

# Numerical Targets and Basic Policy on Profit Distribution



## Numerical Targets

	FY2021 results	FY2025 target
Net sales	¥34.8 billion	¥40.0 billion
Operating income	¥4.4 billion	¥7.0 billion

≪Assumptions≫

- U.S. market introduction of SI-6603 (treatment for lumbar disc herniation)
- Profit expansion from joint function improving agents in Japan
- Expansion of the overseas pharmaceutical and LAL businesses
- Ratio of R&D expenses to sales (excluding royalty income): target of 25%
- Exchange rate: ¥135/US\$1

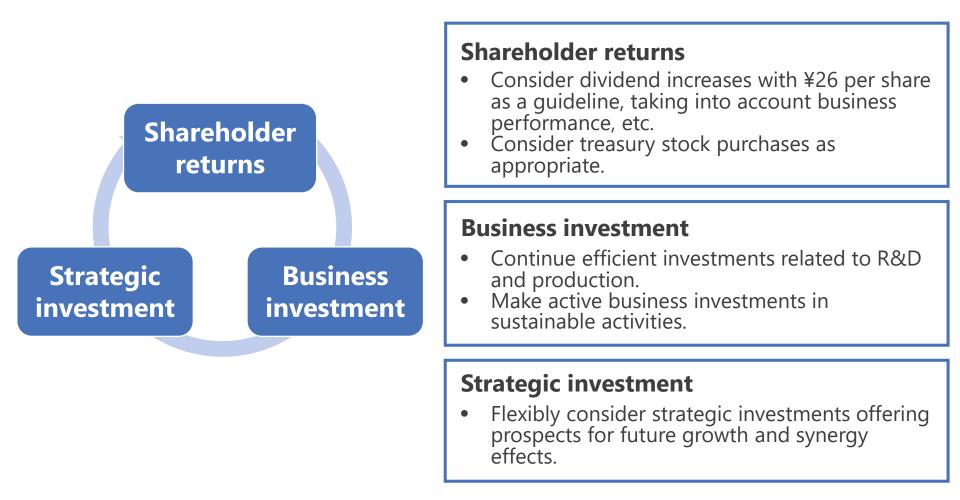


# Aim to achieve record-high business results



Exploring the Innovative Promise of Glycoscience

# Make investments for sustained profit growth and corporate value enhancement and pay stable dividends





# Resolution to revise the dividend in accordance with the dividend policy Now purchasing treasury stock

### FY2022

**Annual dividend : <u>¥26 per share</u>** (including an interim dividend of ¥13) <u>Reference information</u> Previous forecast announced May 13: Annual dividend of ¥22 per share (including ¥11 interim dividend)

	FY2018	FY2019	FY2020	FY2021	FY2022 (forecast)
Net profit per share	¥39.76	-¥192.15	¥75.54	¥66.32	¥48.12
Annual dividend per share	¥26.00	¥26.00	¥24.00	¥30.00 <sup>*2</sup>	¥26.00
Dividend payout ratio	65.4%	-	31.8%	45.2%	54.0%

\*1 Includes JOYCLU approval commemorative dividend of ¥4 \*2 Includes JOYCLU launch special dividend of ¥10

## **Treasury Stock Acquisition**

Total number of shares to be acquired: 2,000,000 shares (maximum) Total value of shares to be acquired: ¥1.5 billion (maximum) Acquisition period: May 16, 2022 to December 30, 2022

#### Totals as of October 31, 2022

Total number of shares acquired: 1,327,800 shares Total amount of acquisition cost: ¥1,127,473,900





#### **Business foundation**

Enhancement of employee engagement/ Organizational strengthening and human resource development

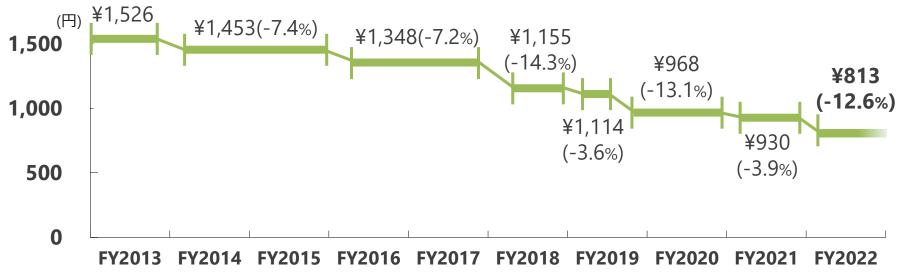
# Nurture the capability to maintain a constant growth trajectory by energetically implementing the key measures

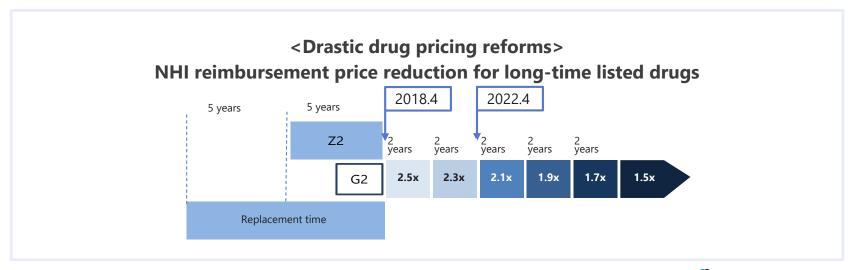


**Exploring the Innovative Promise of Glycoscience** 

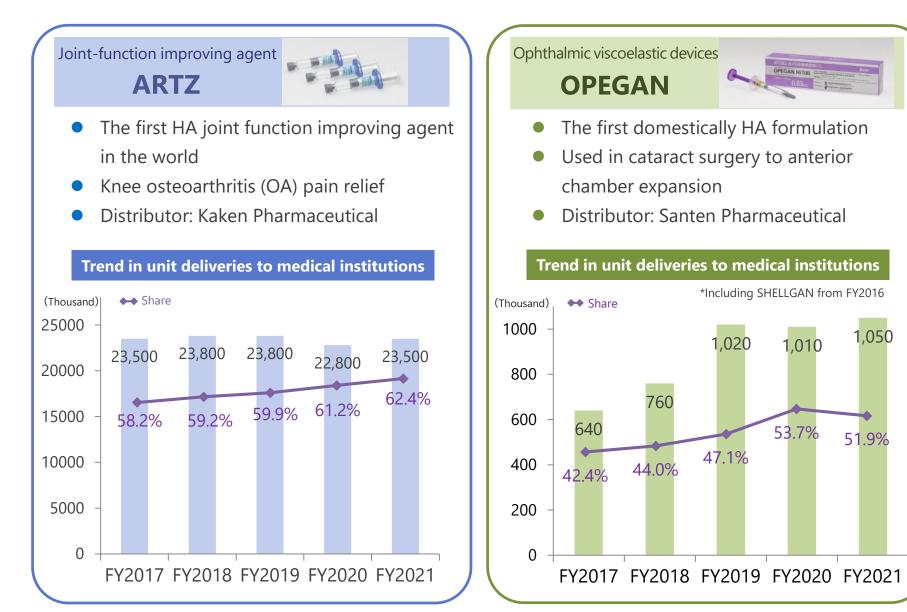








## Trend in Deliveries of ARTZ / OPEGAN to Domestic Medical Institutions





# **Overview of JOYCLU**

# Manufacturing and marketing approval in Japan obtained in March 2021 For the indication of osteoarthritis (knee joint and hip joint) May 19 NHI drug price listing and launch



Product name	JOYCLU® 30mg Intra-articular Injection
Generic name	diclofenac etalhyaluronate sodium
Indication	Osteoarthritis (knee joint and hip joint)
Dosage and administration	The usual adult dosage is 1 syringe per dose (30 mg of diclofenac etalhyaluronate sodium) injected intra-articularly every 4 weeks.



# **Overview of JOYCLU**

Improvement of symptoms expected by administration once every 4 weeks First joint function improvement agent in Japan indicated for the treatment of osteoarthritis of the hip joint

### **Features**

- ► Hyaluronic acid and diclofenac chemically bound using a proprietary technology
- Diclofenac released by hydrolysis in the joint
- Improvement of symptoms of osteoarthritis (knee joint and hip joint) expected by administration once every four weeks
- First joint function improvement agent in Japan indicated for the treatment of osteoarthritis of the hip joint

## **Expected positioning**

Establishment as a new base drug in the treatment of osteoarthritis alongside existing hyaluronic acid formulations and NSAIDs

### <Target Patients>

- > People for whom existing hyaluronic acid formulations are insufficiently effective
- People who want to avoid NSAIDs, such as oral drugs or patches, in consideration of side effects
- ▷ People for whom frequent hospital visits are a hardship
- Provision of a new treatment option for osteoarthritis of the hip joint



# Four types of lumbar disc herniation

#### **1. Protrusion** 2. Subligamentous extrusion Although the hernia extends beyond the The hernia does not extend beyond the outermost layer of the annulus fibrosus, it is outermost layer of the annulus fibrosus covered by the posterior longitudinal ligament Longitudinal section Cross-section Posterior longitudinal Vertebral igament Annulus body fibrosus umbar. disc Nucleus pulposus Posterior longitudinal ligament 3. Transligamentous extrusion 4. Sequestration The hernia perforates the posterior longitudinal The hernia migrates outside the dura mater ligament



# Setting of physician and facility requirements for HERNICORE use to promote appropriate use and ensure safety

#### [Physician requirements]

#### [Japanese Society for Spine Surgery and Related Research]

- Physicians under the following conditions (as of April 2019)
- 1. Supervisory physicians or certified physicians of the Japanese Society for Spine Surgery and Related Research (JSSR) or who are supervised under the JSSR, or who participated in the this clinical study
- 2. Physicians having experience with performing intradiscal puncture or at least 50 surgeries for lumbar disc herniation

#### 【 Neurospinal Society of Japan 】: Physicians under the following conditions (as of April 2019)

- **1.** Supervisory physicians or certified physicians of the Neurospinal Society of Japan
- 2. Physicians having experience with performing intradiscal puncture or at least 50 surgeries for lumbar disc herniation

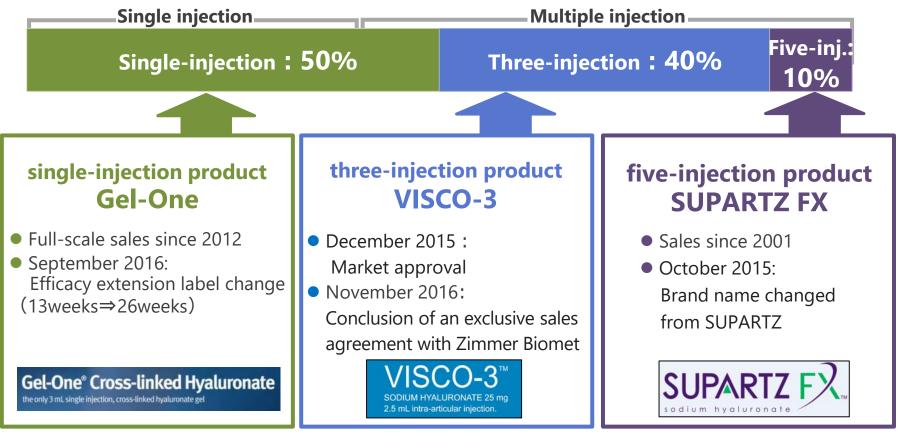
#### **[Facility requirements]** : Facilities under the following conditions

- **1.** Facilities equipped with an X-ray fluoroscopic system (C-arm, etc.) capable of administering HERNICORE using clean technique
- 2. Facilities capable of treating shock and anaphylaxis
- **3.** Facilities capable of performing urgent spine surgery or facilities that cooperate with facilities capable of performing spine surgery
- 4. Facilities with hospitalization equipment
- X Among those who meet the physician requirements, supervisory physicians in the JSSR may be used at non-fulltime work facilities that meet the facilities requirements (as of November 2019)



# Market size of US\$950 mil. in 2021 (+8.3% year-on-year)

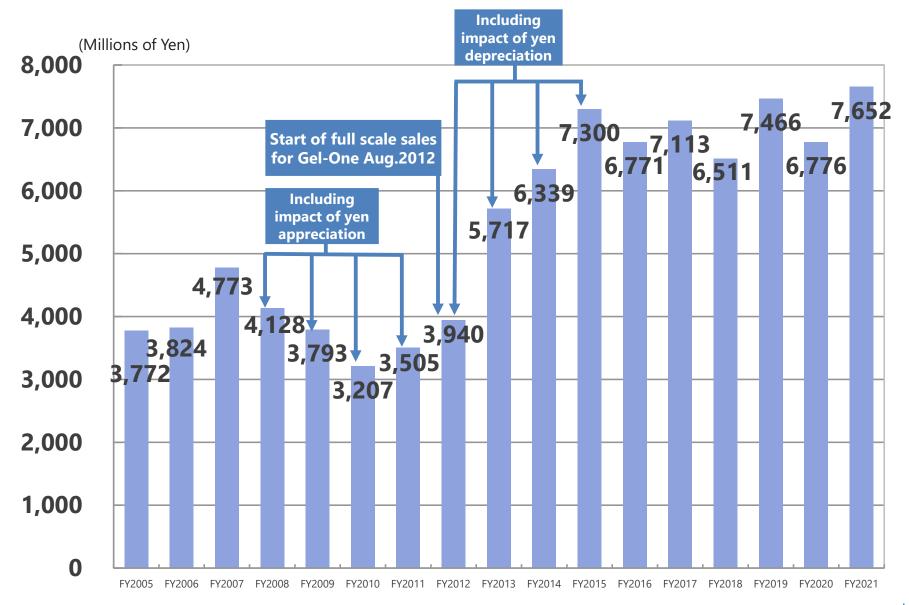
**U.S. market share by number of injections** (Value basis, including competitors)



\*Figures for 2021, Seikagaku estimates



# **Trend in Overseas Sales of Hyaluronic Acid Products**





# **Recombinant LAL reagent** [PyroSmart NextGen]

## April 2021 launch at ACC Product to be marketed globally, with a launch in Japan planned for May or later in 2021



#### **Product Features**

- Product manufactured using recombinant technology without using blood harvested from horseshoe crabs, a raw material used in traditional products
  - $\rightarrow$  Ensures continuous product supply
- Ability to utilize the same test methods and instruments as naturally sourced products
  - $\rightarrow$  Ensures consistency with endotoxin-testing reagents
  - → A next-generation BET reagent designed to deliver highly reliable quantitation of endotoxins



# The LAL Business

## Endotoxin detection reagents (Seikagaku Corporation, ACC)

- Manufacture and sale of reagents for detecting endotoxins\* in the manufacturing process of pharmaceuticals, etc. (Japan, USA, Europe, etc.)
- Used mainly at pharmaceutical companies
- Launch of PyroSmart NextGen, a recombinant endotoxin detection reagent, in April 2021
- Accelerated global expansion of recombinant reagents

\*Since endotoxins exhibit strong pyrogenic activity even in minute amounts, they must be rigorously controlled pursuant to regulations in the manufacture of pharmaceuticals and medical devices.

## ► Global market : Approx. ¥2.5 mil

(Seikagaku estimate / including related equipment)

# Glucan detection in vitro diagnostic agents (ACC)

- Manufacture and sales of in vitro diagnostic agents for measuring the glucan\* concentration in blood for use in the diagnosis of deep fungal infections (USA, Europe, etc.)
- Since symptoms of deep fungal infections can be similar to those of COVID-19, the products are used together with COVID tests as necessary to ensure appropriate diagnosis and treatment
- Used mainly at hospitals and testing centers
- Market expansion into additional countries

\*(1 $\rightarrow$ 3) - $\beta$ -D-glucans are structural components of the cell walls of fungi, as typified by molds and yeasts.



- 受託試験サービス(ACC社)
- Contract testing for endotoxin and glucan contamination
- Contributes to quality control at pharmaceutical companies and medical institutions



# Developed the world's first endotoxin detection reagent and obtained FDA approval in 1977 / Sales network spanning 80 countries

## Associates of Cape Cod, Inc. (ACC社)

- Headquarters : Massachusetts, U.S.A.
- Established : 1974 (became a Seikagaku subsidiary in 1997)
- Business areas : Manufacture and sale of endotoxin detection reagents and glucan detection in vitro diagnostic agents, provision of contract test services
- Number of employees : 263 (as of March 31, 2022)



Recombinant LAL reagent PyroSmart NextGen<sup>®</sup>



Endotoxin detection reagents PYROCHROME® Exterior of the ACC office



# Dalton, a Canadian CDMO, becomes a subsidiary Acquiring a pharmaceuticals manufacturing base compliant with US/Canadian GMP

## **Outline of acquisition**

- Name : Dalton Chemical Laboratories, Inc.
- Acquisition cost : Up to CAD 41 million
- Acquisition method : Acquired all issued shares

## **Dalton Chemical Laboratories, Inc.**

- Location : Toronto, Ontario Province, Canada
- Established : 1986
- Business description :

Contract manufacturing services (CDMO\*), including the manufacturing of chemosynthesis products and active pharmaceutical ingredients (API) and API process development for pharmaceutical companies

Employee number : 147 (2022.3)

\* CDMO: Contract Development and Manufacturing Organization A business that supplies comprehensive services in drug development and manufacturing to pharmaceutical companies, including contract drug manufacturing, pharmaceutical formulation planning at the development stage, manufacturing of investigational drugs, and optimization of manufacturing conditions.





# Established new company in Canada, promoting development of Seikagaku pharmaceuticals and medical devices in North America

## SEIKAGAKU NORTH AMERICA CORPORATION

- Location : Toronto, Ontario Province, Canada
- Established : January 25, 2022
- Capital : CAD10
- Business description : Development of pharmaceuticals and medical devices in North America

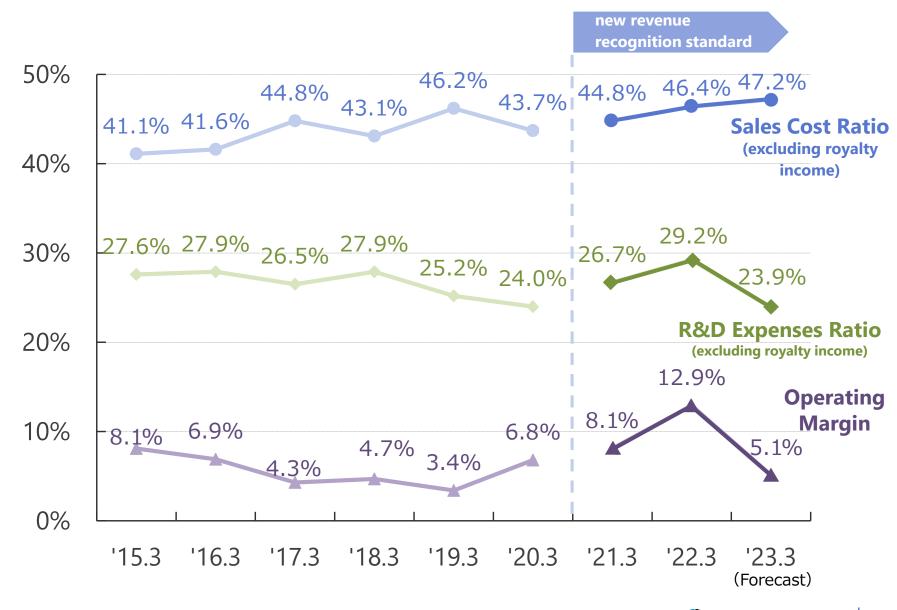
## **Purpose of establishment**

- Enable responses with fewer constraints on time differences or distance by operating a development base in North America
  - Formulate development plans more closely aligned with local medical environment
  - Create smooth communications with U.S. Food and Drug Administration (FDA) and clinical trial sites

Strengthening the development system in North America. Aiming to accelerate pharmaceutical and medical device development as well as obtaining of approvals

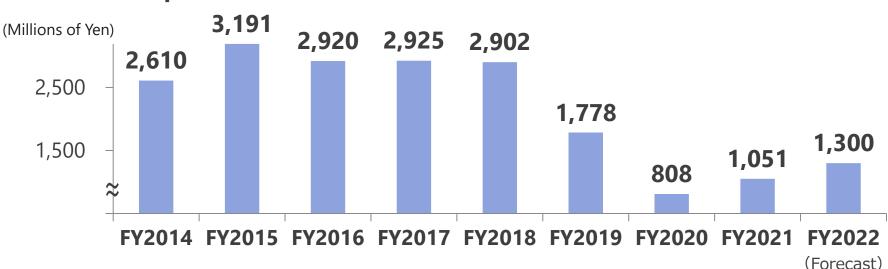


# **Trend in Financial Index**



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# Impairment loss taken in FY2019 Depreciation will trend toward ¥1.0 billion



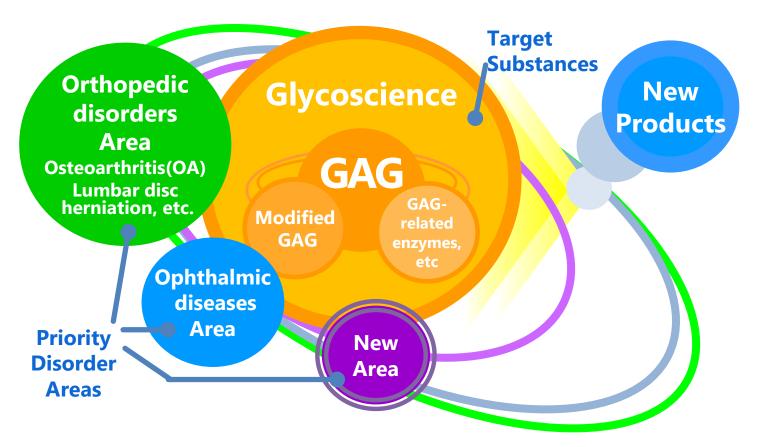
## Trend in Depreciation

**Trend in Capital Investments** (Millions of Yen)

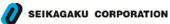
FY2014	FY2015	FY2016	FY2017	FY2018	FY2019	FY2020	FY2021	FY2022 (Forecast)
2,095	1,975	1,173	1,591	1,310	2,109	2,127	2,194	3,000



# Continue Our R&D policy to make the most of our technology and knowledge



GAG: Glycosaminoglycans (One of the constituents of complex carbohydrates)



# Development of SI-613-ETP discontinued Prioritize identification of cause of JOYCLU side effects



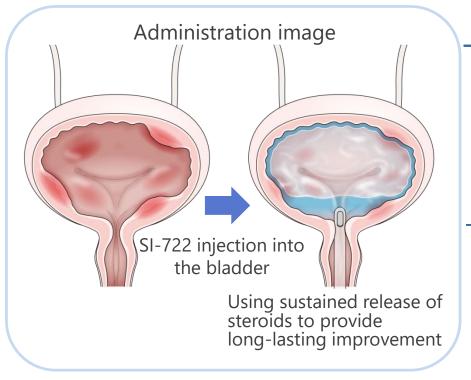
### **Development status**

- SI-613 (osteoarthritis of the knee)
   U.S. /China/South Korea
  - Proceeding with preparation of a clinical development plan
- SI-613 (Enthesopathy) Japan
  - February 2022: Development discontinued
    - ⇒ Failure of the PII clinical study to meet its primary efficacy evaluation, and to prioritize identification of the cause of shock or anaphylaxis occurring in patients following administration of JOYCLU

<SI-613 summary>
Dev. code : SI-613 Generic name : Diclofenac etalhyaluronate sodium
Indication : Osteoarthritis/Enthesopathy Method of use : Injection into joint cavity
Estimated patients : 24 million (U.S.) / 47 million (China) 3.7 million (South Korea)

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# Subject enrollment for PI/PII studies in the U.S. completed Analysis of study results to proceed



## **Development status**

- ► U.S. Phase I/II : Starting November 2019
  - Subject enrollment complete in January 2021
  - ⇒Tolerability confirmed (a primary objective) Will consider the next-phase study

## **Expected Features**

- Designed for sustained release by bonding steroids to chondroitin sulfate
- Anti-inflammatory effects of steroids under sustained release should provide sustainable relief of frequent urination and bladder pain

<SI-722 summary>

Dev. Code: SI-722Generic name: Steroid conjugated with chondroitin sulfate Indication: Interstitial cystitis and bladder pain syndrome Method of use : Injection into the bladder Estimated U.S. patients : 1.3 million (Seikagaku estimates) SEIKAGAKU CORPORATION



# **Clinical Study Information**

Development code/ Indication	Develop- ment Location	Clinical Study Title (Study ID)	Target Enroll- ment	Estimated Period	Primary End Point (Primary Follow-up period)
<b>SI-6603</b> Lumbar Disk Herniation	U.S.	Phase III additional study ( <u>NCT03607838</u> )	320	May. 2018 – Nov. 2022	Leg pain (13 weeks)
<b>SI-613-ETP</b> Enthesopathy	Japan	Late-stage Phase II clinical study (JapicCTI-173758)	240	Oct. 2017 – Oct. 2018	Pain in motion (4 weeks)
<b>SI-613</b> Knee Osteoarthritis	U.S.	Phase II clinical study ( <u>NCT03209362</u> )	80	Oct. 2017 – Nov. 2018	WOMAC(Knee pain) (12 weeks)
<b>SI-614</b> Dry eye	U.S.	Phase III clinical study	240	May. 2022 – Jan. 2023	
<b>SI-722</b> Interstitial cystitis and bladder pain syndrome	U.S.	Phase I / II clinical study ( <u>NCT04208087</u> )	32	Mar. 2020 – Jan. 2021	Maximum observed plasma concentration
<b>SI-449</b> Adhesion Barrier	Japan	Pivotal study (Field of gastroenterological surgery) (JapicCTI-205343)	130	Jun. 2020 – Dec. 2022	Efficacy
<b>SI-449</b> Adhesion Barrier	Japan	Pilot study (Field of gynecology) (jRCT2072210100)	10	Dec. 2021 – Sep. 2022	Usability, Safety

Note: The table shows data registered (or planned to be registered) on clinical trial information websites.

The information is updated from time to time. Refer to the websites for details and the latest information. (The websites can be accessed from the trial ID links.)

• Japan Pharmaceutical Information Center(JAPIC) <u>http://www.clinicaltrials.jp/user/cteSearch\_e.jsp</u>

University hospital Medical Information Network (UMIN) Center http://www.umin.ac.jp/ctr/index.htm

ClinicalTrials.gov <u>https://clinicaltrials.gov/ct2/search</u>

Note: Actual enrollments or trial periods may differ from targets and plans due to various factors.



# Planned receipt of milestone royalties in accordance with future progress in development and marketing

Development Code Indication	Develop- ment Location	Sales Partner	Total Amount of Milestone Royalties ( of which, in upfront payment )
<b>SI-6603</b> Lumbar Disk Herniation	U.S.	Ferring Pharmaceuticals (Switzerland)	Max. US \$95 million (US \$5 million)
	Japan	Ono Pharmaceutical Co., Ltd.	Max. ¥12.0 billion (¥2.0 billion)
SI-613	U.S.	Searching	—
Japan: Osteoarthritis U.S. : Knee Osteoarthritis	China	Eisai Co., Ltd.	_
	Korea	Eisai Co., Ltd.	—
SI-613-ETP Enthesopathy	Japan	Ono Pharmaceutical Co., Ltd.	*included in the above
SI-614 Dry eye	U.S.	Searching	—
SI-722 Interstitial cystitis	U.S.		—
SI-449 Adhesion Barrier	Japan	—	_





A company that is valued by the world through its innovative drug discovery
Creativity, Fairness, Dreams and Passion
We create safe and useful products for human well-being with basic research based on glycoscience.
<ul> <li>We create a corporate environment of mutual trust and communication using individual abilities.</li> <li>We create innovative and useful products through in-depth cooperation between industrial and academic circles.</li> <li>We assure the highest quality and safety of our products.</li> <li>We enhance interaction with society by establishing genuine trust. Through these efforts, Seikagaku will strive to become a sound and socially responsible company that protects the natural environment and improves quality of life.</li> </ul>
"Glycoscience for human well-being"







# **Specialization in Glycoscience**

- Niche field, market not big enough for the major pharmaceutical companies to penetrate
- Focusing on this field more than 70 years

# State-of-the-art technology related to GAG



 Drug discovery expertise using modified-GAG, GAG-related enzymes, etc.
 Extraction, Purification, Fermentation, etc. technology to manufacture GAG related products



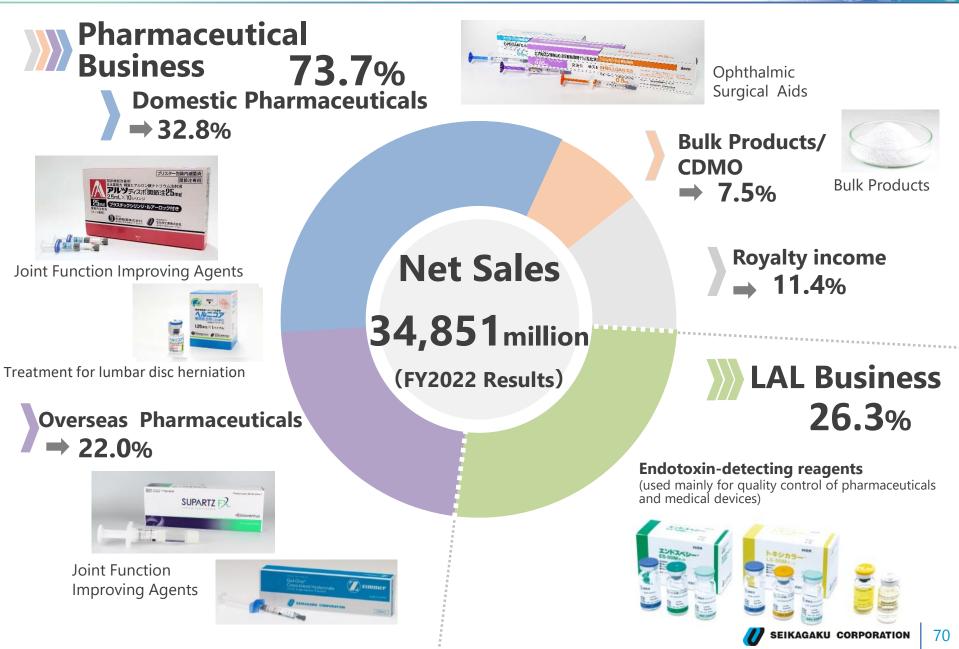
# <u>Unique business model</u>

Concentration on **R&D** and **manufacturing** 

R&D staff comprising 40% of our employees (Non-consolidated base)
 Allocation of 25% of net sales to R&D investment



# **Our Business Segment**



# **ARTZ<sup>®</sup>** Joint function improving agent by multiple injections

- The first HA joint function improving agent in the world
- Main distributors:

Kaken Pharmaceutical (Japan): ARTZ

Bioventus (U.S.): SUPARTZ FX

Kunming Baker Norton

Pharmaceutical (China): ARTZ



# **Gel-One**<sup>®</sup> Intra-articular single-injection viscosupplement for the treatment of knee osteoarthritis

- Requires only 3ml to be as effective as multiple injections
- Full-scale sales since August 2012
- Distributor: Zimmer (U.S.)





#### ■ Cautionary Notes

This material contains forward-looking statements regarding future management strategies or performance forecasts. These descriptions are based on judgments derived from information that is currently available to Seikagaku and are subject to risk and uncertainty. Actual results and developments may differ significantly from these descriptions due to various factors. Information about pharmaceutical products or medical devices (including products currently in development) contained in this material is not intended to constitute an advertisement or medical advice.



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(TSE: 4548)

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