





# Bio CDMO Div. & Life Sciences Business Div. Business Briefing

# **FUJIFILM Holdings Corporation**

#### January 6, 2022

Forward-looking statements, such as those relating to earnings forecasts and other projections contained in this material, are management's current assumptions and beliefs based on currently available information. Such forward-looking statements are subject to a number of risks, uncertainties, and other factors. Accordingly, actual results may differ materially from those projected due to various factors.

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# Opening & Introduction : Organization and Mid-term Plan for Healthcare Takatoshi Ishikawa

FUJIFILM Corporation Director, Senior Executive Vice President, CLSO General Manager, Bio CDMO Division

# Bio CDMO Division Takatoshi Ishikawa

FUJIFILM Corporation Director, Senior Executive Vice President, CLSO General Manager, Bio CDMO Division

# Life Sciences Business Division Yutaka Yamaguchi

FUJIFILM Corporation General Manager, Life Sciences Business Division



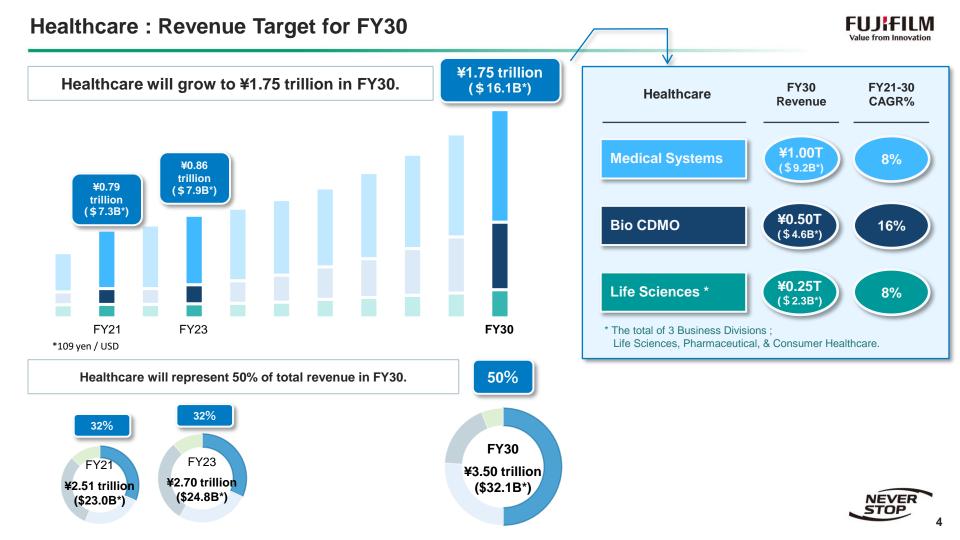
#### Healthcare / Life Sciences : New Organization (Effective on April 1, 2021)



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- Redefine "Healthcare" as two business groups ; "Medical Systems" and "Life Sciences".
- Reorganize and strengthen Life Sciences businesses from a customer perspective, to prioritize Bio CDMO and drug development support businesses.
- Become a leader within the life science industry, by offering value of end-to-end solutions as a company strongly supporting the creation of cutting-edge medicine.

Previously (~March 31, 2021)	$\triangleright$	Currently (April 1, 2021~)				
Healthcare		Healthcare				
Medical Systems Business Div.		Medical Systems Business Div.				
Bio CDMO Div.		Life Sciences Bio CDMO Div.	* Page5-			
Regenerative Medicine Div. (Cell / Cell culture media)		Life Science Business Div. Life Sciences (Cell/Cell culture media/Reagent)	* Page28-			
Pharmaceutical Products Div.		Strategy Headquarters Pharmaceutical Products Div.				
Life Science Products Div. (Cosmetics / Supplements)		Consumer Healthcare Business Div. (Cosmetics / Supplements)	NEVER STOP			









# **Bio CDMO Division**

FUJIFILM Corporation Director, Senior Executive Vice President, CLSO General Manager, Bio CDMO Division Takatoshi Ishikawa

January 6, 2022



- 1. Overview of Fujifilm's Bio CDMO Business
- 2. Outlook for Bio CDMO Business
- Advantage of Fujifilm's Technology
   3-1 Technological Differentiation within a Wide Variety of Modalities
   3-2 Industry's First Continuous Production System from Culture to Purification
- 4. Summary



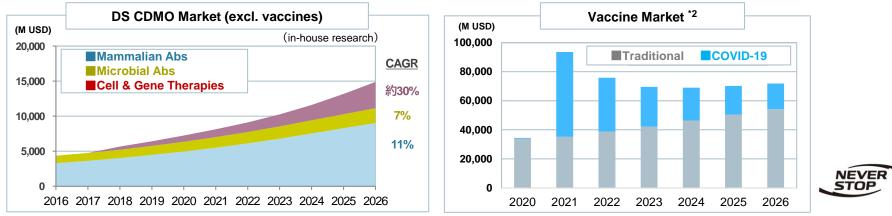
# The demand for CDMOs will increase further due to conventional antibodies, progress within next gen biopharmaceuticals and increased vaccine demand.

 The growth of the Bio CDMO drug substance market is accelerating further and <u>is expected to reach ¥1.3T by 2025 (+¥300B compared to the</u> previous forecast, excl. vaccine demand). The primary changes are:

**1)** Acceleration of cell & gene therapy development : The market CAGR is expected to increase to approx. 30% (+10% compared to the previous forecast). Both the commercial and clinical pipeline have increased (clinical trials in 2020: 1,220 $\rightarrow$ 2021: 1,320<sup>\*1</sup>), and the market is expected to expand even further beyond 2025.

**2)** Increase in demand for next gen biopharmaceuticals: The mammalian cell culture segment is expected to continue to grow at a high CAGR of 11% (+0.5% compared to the previous forecast). In addition to the strong growth of conventional antibodies, new modalities such as the highly effective ADCs (Antibody Drug Conjugates) and bispecific antibodies contribute to the growth.

In the vaccine market, there has been a major increase in demand related to COVID-19 vaccines (2021 market size: ~¥6T\*2). This demand is
expected to continue thereby increasing pressure on the manufacturing capacity of drug substance CDMOs.



#### Fujifilm can handle various modalities, small- to large-scale manufacturing and provide end-to-end solutions in response to the needs of its customers

(In-house market research)

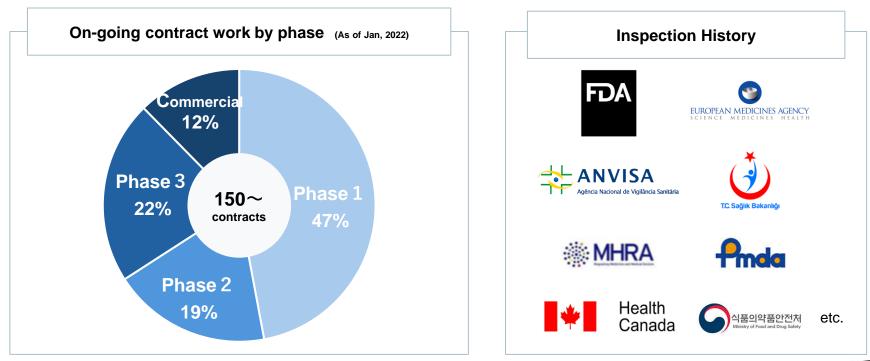
				Fujifilm	Firm A	Firm B	Firm C	Firm D
	Drug Substances	rProtein drugs (Microbial)		~	~	~	-	~
		rProtein vaccines (Insect cells)		~	-	-	-	-
Dru		Gene therapies		<ul> <li>✓</li> </ul>	~	~	-	-
g Sul		Cell therapies		<ul> <li>✓</li> </ul>	~	~	-	-
bstan		Antibodies (mammalian)	Large Scale	~	~	-	~	~
ICes			Small/mid Scale	~	<b>~</b>	~	-	~
	Hybrid	mRNA vaccines		LNP <sup>*</sup> Formulation	~	-	~	-
	Chemical	Small molecule drugs		<ul> <li>✓</li> </ul>	~	~	-	-
	Formulation			~	~	~	~	~

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Within the Europe and U.S., Fujifilm's primary market, the strengths of individual sites are leveraged to handle process development and manufacturing of drug products from clinical to commercial stage products

*As of Jan 2022					New	New
	Billingham UK	North Carolina US	Texas US	Hillerød Denmark	Boston US (operational)	North Carolina US (TBO 2025)
				and the second s		
Services						
Antibodies	$\checkmark$	$\checkmark$	$\checkmark$	$\checkmark$		$\checkmark$
Recombinant protein	$\checkmark$	$\checkmark$				
Cell/gene therapies	$\checkmark$		$\checkmark$		$\checkmark$	
Vaccines	$\checkmark$	$\checkmark$	$\checkmark$			
Formulation			$\checkmark$	$\checkmark$		$\checkmark$
Assembly & packaging				$\checkmark$		$\checkmark$
FUJ:FILN Diesyntl biotechnologie			<b>FUJ:FILM</b> Fujifilm HQ okyo, Japan)	o 9º FL	JJ:FILM Diesynth biotechnologies	

# Achieve continuous growth by contracting a large amount of early clinical stage (Phase 1, 2) projects with future potential







## 1. Overview of Fujifilm's Bio CDMO Business

## 2. Outlook for Bio CDMO Business

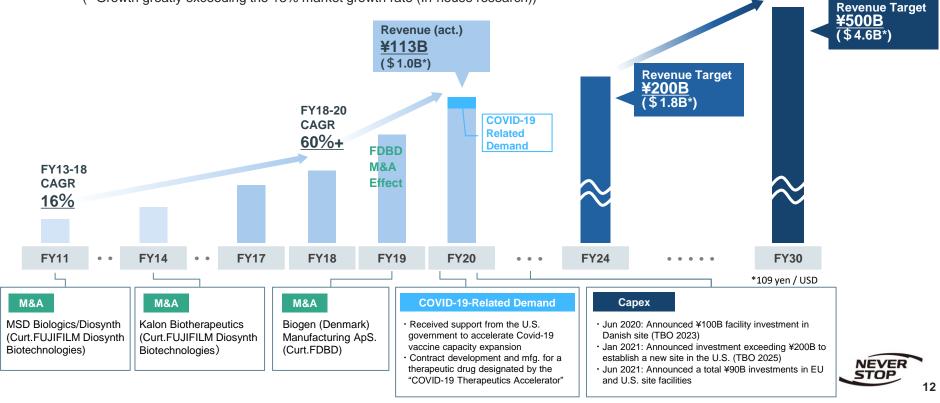
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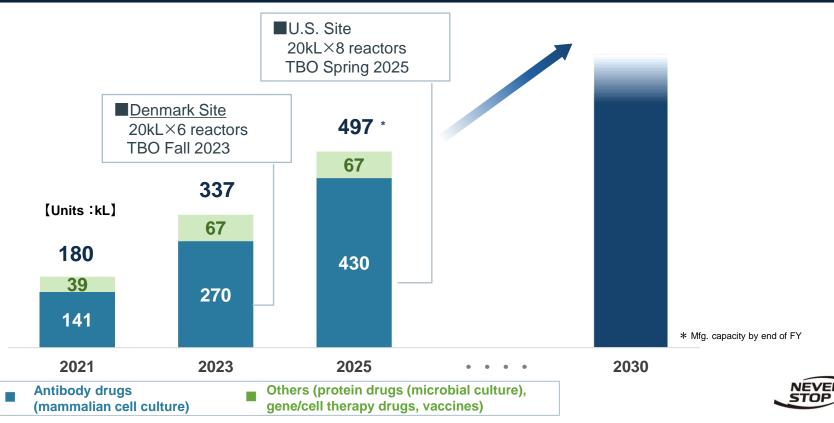
Will expand business through active facility investments and M&A to aim for a revenue of ¥200B in 2024 and ¥500B in 2030

(\* Growth greatly exceeding the 13% market growth rate (In-house research))





Will expand manufacturing capacity for all modalities at a rate exceeding the market growth rate to achieve rapid growth





#### Large-scale expansion of drug substance, fill & finish, and packing facilities underway at Fujifilm's site in Denmark. \* \* Largest scale investment in Denmark Ttl floor area : $40,000 \text{ m}^2 \rightarrow 60,000 \text{ m}^2$ | Ttl tank volume : $120 \text{ kL} \rightarrow 240 \text{ kL}$ • 20kl Bio reactors Current Status (Nov 2021) Curt. $6 \rightarrow 12$ On-going Expansion • New building with all-automated fill & finish system Expansion of assembly, labeling, packaging facilities \*Denmark Site : - Site area 250,000m<sup>2</sup> (x35 soccer fields) Further room for expansion - Aim to achieve carbon neutrality by 2025 2021 2022 2023







New site under construction in North Carolina with <u>20,000 x 8 bio reactors, fill & finish and packaging</u> <u>capabilities</u>. Set to become the largest\* bio CDMO site in Northern America.

- 20kl Bio reactors
- $8_{\text{construction}} \rightarrow \text{Maximum } 32_{\text{expanded}}^{\text{Can be}}$
- All-Automated fill & finish system
- Assembly, labelling, packaging

\*New NC, U.S. Site :

- Site area 610km (x85 soccer fields)
  - ➡ The largest\* bio CDMO site in Northern America
- Plan to use 100% electricity from renewable sources







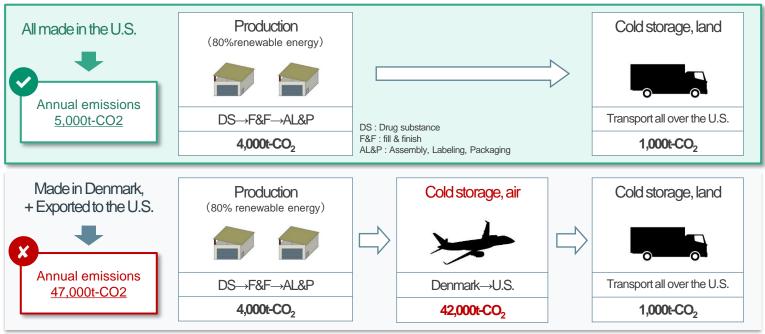
\*Viewed by +10,000 people on social media



# Fujifilm aims to reduce the environmental burden of the pharmaceutical supply chain by promoting "local production for local consumption".

#### Relative amount of CO2 emitted (In-house simulation\*)

\* [Premise] Relative CO2 emissions before reaching patients when mfg. the same amount of the same drug for the U.S. market



Significant amount of CO2 emissions when manufacturing products for the U.S. within the EU due to the need for cold storage and air transport



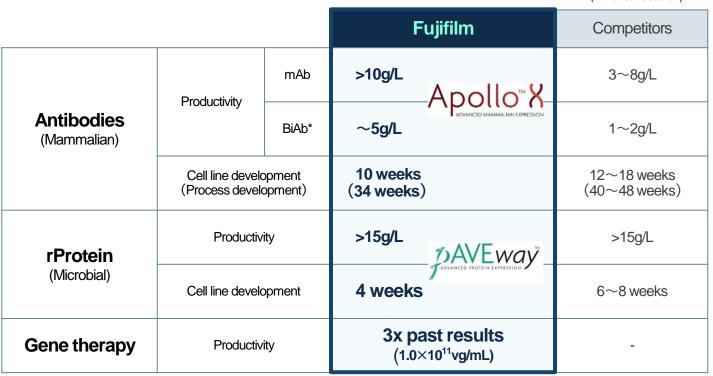


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#### Industry-leading productivity for all modalities



(in-house research)

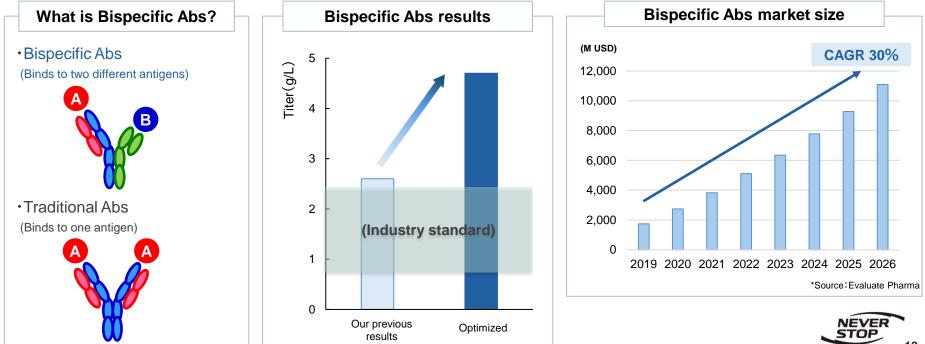


\*BiAb: bispecific antibodies

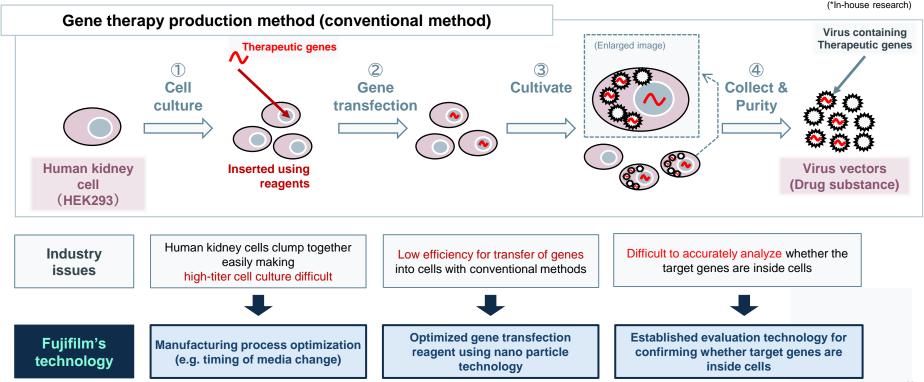
Bio CDMO Div FUJIFILM Value from Innovation

Fujifilm has begun contract manufacturing of bispecific antibodies and using proprietary plasmid transfection technology has resulted in the creation of a cell line with <u>industry-leading productivity (~5g/L)</u> \*

(\*Announced at the "Antibody Engineering & Therapeutics" conference on Dec. 15th, 2021)



Fujifilm has, as a CDMO, established its <u>own original manufacturing process</u> thereby <u>tripling drug substance productivity</u> through process technology, nano particle control technology and AI analysis.





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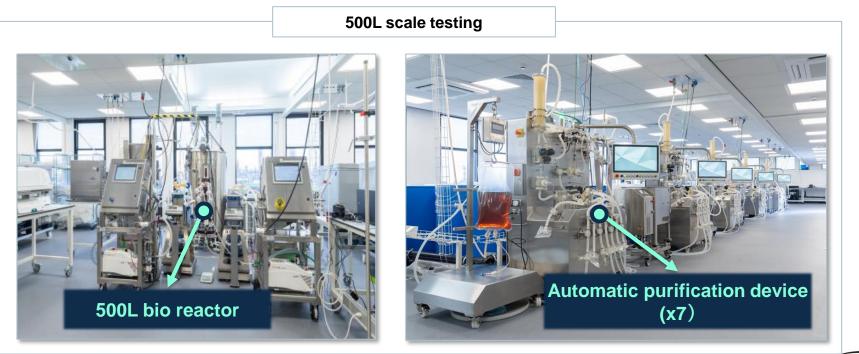
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#### **3-2-1 | Continuous Manufacturing from Culture to Purification**



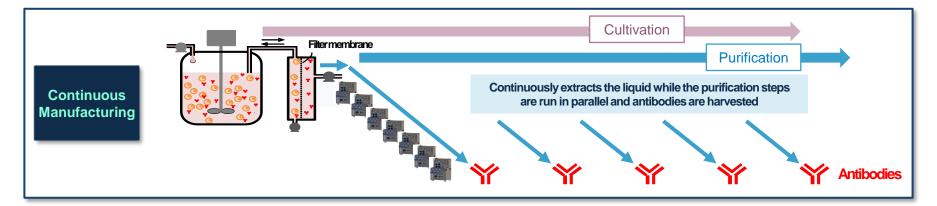
# Have developed the world's first 500L scale continuous manufacturing facility and <u>have gathered empirical evidence</u>. Have also developed original media for each piece of equipment.

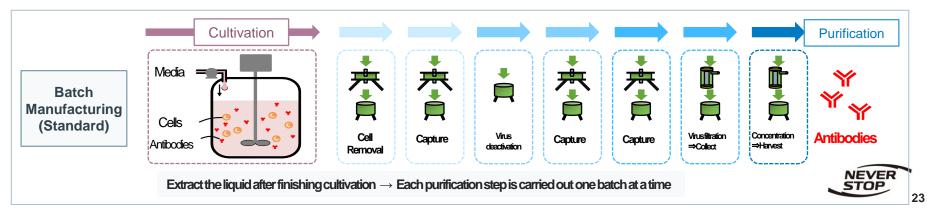






#### Continuously extracts the liquid while the purification steps are run in parallel and antibodies are harvested





#### Set to begin GMP operation from 2023 and are currently having discussions with several clients

	Features of Fujifilm's Continuous Manufacturing	Batch Production
Quality	<ul> <li>Ability to achieve high purity compared to batch production</li> <li>Enables manufacturing of unstable antibodies that are difficult to produce with batch production.</li> </ul>	<ul> <li>Unstable antibodies are hard to manufacture</li> </ul>
Production capacity	<ul> <li>By adjusting the production time small to large scale lots can be made at the same facility</li> </ul>	Different facilities needed for different lot sizes
Facility investment Mfg. cost	<ul> <li>Takes up 25-75% less space compared to batch production</li> <li>Facility investment amount is likewise reduced by 25-75%</li> <li>25% reduction in manufacturing costs(In-house research)</li> </ul>	<ul> <li>Need to invest in bio reactors depending on the amount to be manufactured</li> </ul>
Technology	<ul> <li>Systems for automatic titer control and continuous monitoring of culture conditions are necessary (development complete)</li> <li>The automatic continuous manufacturing device also needs an automatic control system (development complete)</li> </ul>	-
Culture media	<ul> <li>Media optimized for continuous manufacturing is necessary and Fujifilm has developed a high-quality media for this purpose.</li> </ul>	-



NEVER



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# **1**Strengthen offering for various modalities<br/> $\rightarrow$ Especially for next gen therapies (e.g. ADCs and cell therapies)

2 Pursue high productivity for each modality → Strive to become the industry leader not only within Antibody drugs and recombinant protein drugs, but also cell and gene therapies.

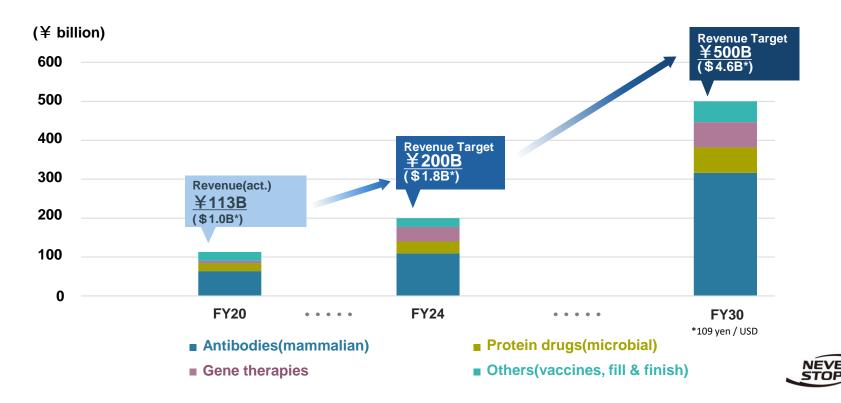
3 Continue timely facility investments → Currently considering a hybrid facility investment of batch and continuous manufacturing.



4 Environmental awareness
 → Localization of production with U.S. site and use of renewable energy to reduce the environmental burden.



Developing industry leading technology and increasing manufacturing capacity allows us to aim for an annual revenue of <u>¥200B in 2024</u> and <u>¥500B in 2030</u>









# **Life Sciences Business Division**

FUJIFILM Corporation General Manager, Life Sciences Business Division Yutaka Yamaguchi

January 6, 2022



April 1990	Joined Fuji Photo Film Co., Ltd. and assigned to the Exportation Division
March 1994	Fuji Photo Film Singapore Office
July 2007	FUJIFILM Electronic Imaging Europe GmbH
April 2014	GM, Life Science Products Division (current Consumer Healthcare Business Division)
May 2018	FUJIFILM Holdings America Corporation, GM, Life Science Strategic Business Office(current position)
June 2018	FUJIFILM Irvine Scientific Inc. CEO (current position)
April 2021	<b>GM, Life Sciences Business Division (current position)</b> Director, The Forum for Innovative Regenerative Medicine (FIRM)





- 1. Positioning and Locations of Fujifilm's Life Sciences Business
- 2. Drug Discovery & Manufacturing Support
- 3. Cell Therapy Process Development & Manufacturing Service
- 4. Summary



#### Handling R&D and production of innovative drugs, and offering solutions in the field of cell therapies to contribute to addressing unmet medical needs

**Drug Discovery & Manufacturing Support** 

- Supplying cells (e.g. human iPSCs for drug discovery), culture media, reagents and related products to contribute to discovery research and production of new innovative drugs.
- Cell culture media has grown rapidly due to increased demand for use in the manufacturing of biopharmaceuticals including COVID-19 vaccines and therapeutics.



Cell Therapy Process Development & Manufacturing Service

 Promoting efficient R&D, with a focus on FCDI's<sup>\*</sup> iPSCs, through Fujifilm group-wide synergy of unique engineering technologies /resources, and alliance with partners.

\* FCDI : Fujifilm Cellular Dynamics, Inc

 Using Fujifilm Group companies' technological resources and facilities as a platform for promoting the process development and manufacturing business for cell therapy products.



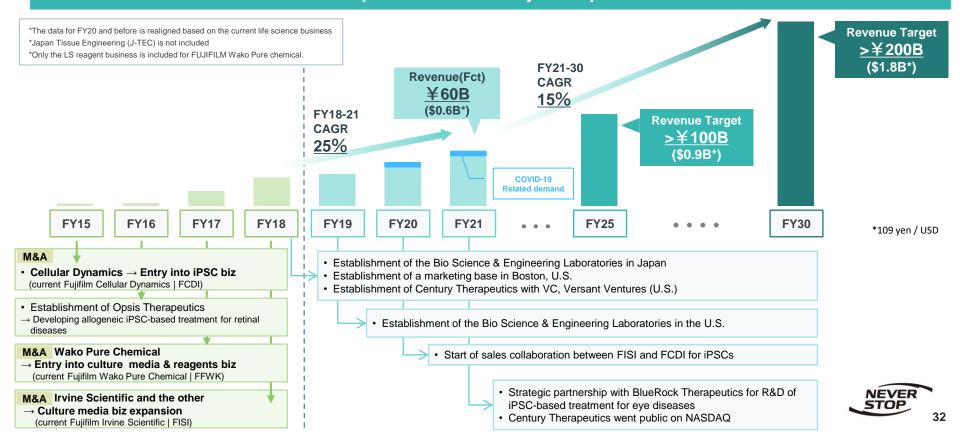
**GMP facility : i-FACT** (Madison, Wisconsin, USA)





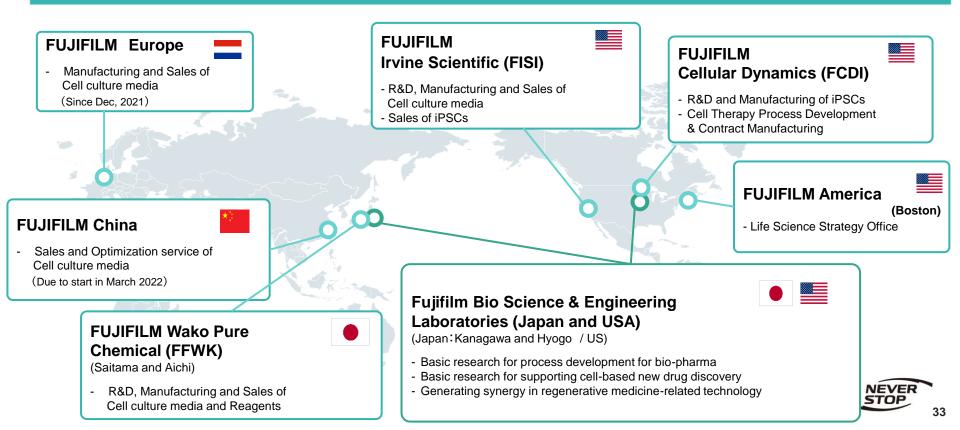
#### 1-2 | Start-up and Expansion of Life Science Business

# Aim to reach $\pm 200B$ in FY2030 by implementing strategic measures one after another since 2018 upon acquisition of three major companies



### 1-3 | Global Network

Generating synergy between Group companies, using the strength of having a wide range of technologies associated with cells, cell culture media and reagents





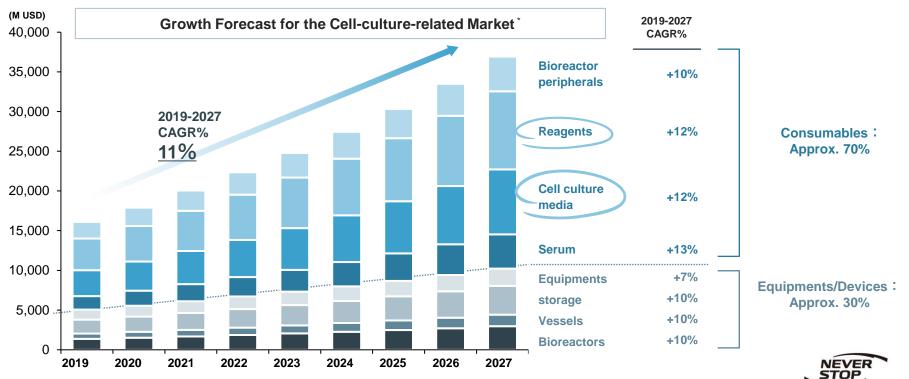
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The cell culture-related market is predicted to grow from \$16.1B in 2019 to \$36.9B in 2027 (CAGR+11%). Consumables such as cell culture media and reagents, which represent Fujifilm's main business category, account for 70% of the market.



\*Source: In-house research based on the report of "Allied Market Research (published in December 2020)"

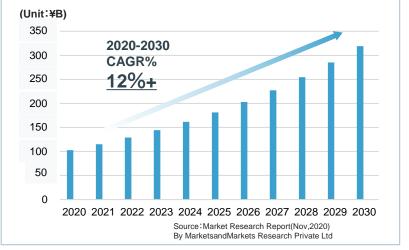
#### Bioproduction media Fujifilm is focusing on is the growing market with CAGR +12%. Fujifilm will be a market leader for BP Media holding >30% market share with ¥100B sales by 2030.

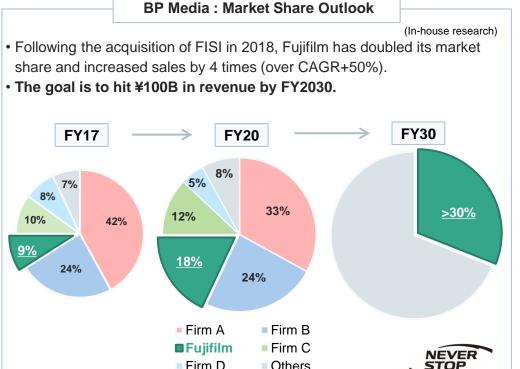
**BP Media : Global Market Outlook** 

 Total demand of serum-free culture media<sup>\*</sup> for bioproduction (BP), a focus area for Fujifilm, is expected to grow at the rate of CAGR+12%.

\*Serum-free culture media:

Serum-free culture media is formulated to mitigate risks from the use of undefined and highly-variable serum products. For this reason, serum-free culture media are widely used for commercial production for biomedicine."





#### Coordinating departments to provide powerful customer support to trade with 15 out of the world's top 20 pharmaceutical companies

R&D	Manufacturing	Sales & Operation
<ul> <li>More than 50 years of cell culture media expertise and an advanced media portfolio.</li> </ul>	Using GMP-compliant manufacturing facilities to produce high-quality culture media.	• Deploying an enhanced product line-up including culture media for broad-based applications, buffers and sterile water for
Ability to provide solutions by		injection.
conducting quick testing at FCDI+FDB and	Products-supply from a global	
optimizing through "cells / culture media /	production framework consisting of sites	Robust global sales networks, combined
culture processes".	in Japan, EU and USA.	with sales teams with outstanding expertise.
Applying the state-of-the-art technology	Using advanced analysis technology, etc.	
in powder and liquid process, developed through the photo film business.	for Quality Control & Quality Assurance.	

Handling processes from development to manufacturing and quality assurance swiftly to supply high-quality products in a timely fashion

Identifying customer needs accurately and providing it to R&D as feedback

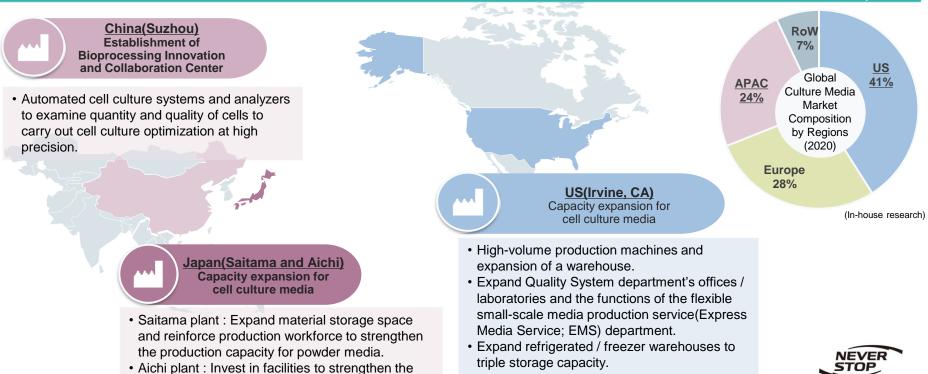


## 2-4-1 | Cell Culture Media : Strategic Investment(US, Japan, China)

supply of process solutions for vaccine production.

Life Science Biz Div Value from Innovation

Cumulative capex of ¥11B (\$100M\*) since acquiring FISI increased production capacity by 5 times.
 Capacity expansion (US, Japan) and establishment of customize service site (China) were decided in FY21.
 Investments will continue into FY2022 and beyond.



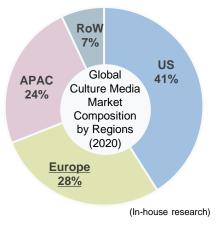
Set within FUJIFILM Manufacturing Europe B.V. the world-class cell culture media facility acting as a hub for the European market.

Enhance global production structure with modern manufacturing sites in Japan, the US and Europe.

Netherlands(Tilburg) Operation commencement (Dec 2021)

- Overview of the new facility
- Total floor space: 13,500m (145,300ft<sup>2</sup>)
- Production Capacity : Dry powder media 320t/yr, Liquid media 470kl/yr
- **Diverted** the existing photographic film plant with high-precision manufacturing experience in Netherlands to culture media plant
- Carbon neutrality by the end of FY2022



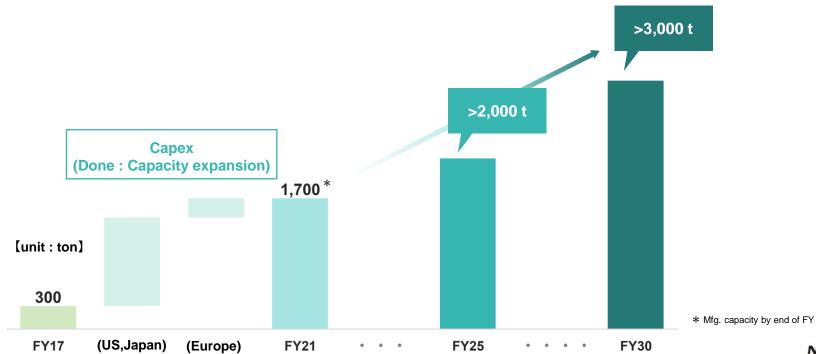






Life Science Biz Div

# Continuous capex worldwide will rise the production capacity in 2030 tenfold compared to FY17 (acquisition of FISI) .

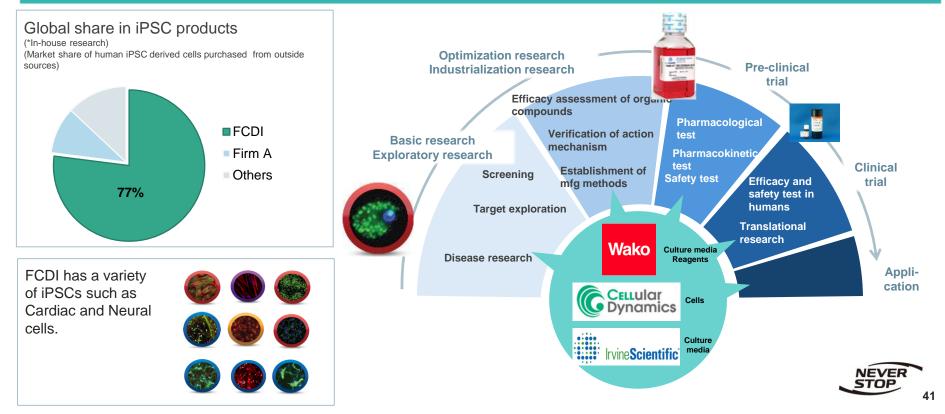




## 2-6 | Cells/Culture media/Reagents : Business Expansion Strategy



Take advantage of cells, culture media and reagents to support drug discovery activities by pharmaceutical companies and academia Engage in drug development projects with customers from an early stage as a "best partner"



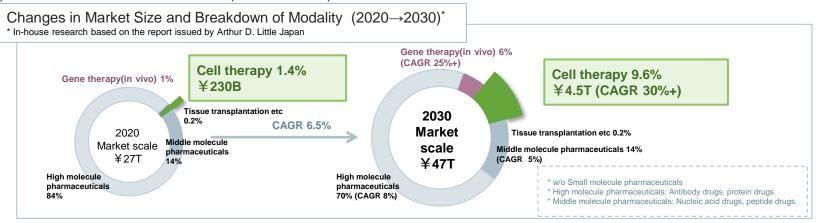


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Cell therapy market has high future growth potential of CAGR 30%+ In US, Investment to biotech companies dedicating to the development of iPSC derived Cell therapies is becoming active, resulting in over ¥1T investment.

- Outlook of the Global Cell Therapy Market (w/ ex-vivo gene therapies)
  - 1) Market size will expand through the shift of categories from autologous to allogeneic, and from somatic / somatic stem cells to iPS cells / ES cells 2)  $\pm 230B$  in 2020  $\rightarrow \pm 4.5T$  in 2030<sup>\*</sup> (CAGR +30%+) \* Source: BB Bridge "FY2020 / Current status and future outlook of cell pharmaceutical development in the world



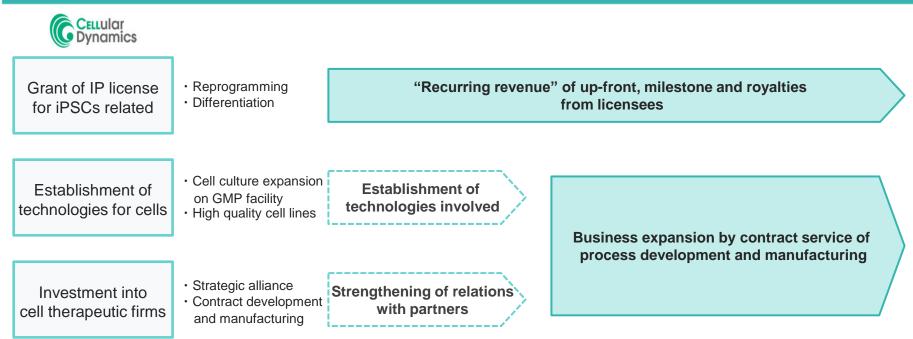
• Corporate values dedicating to the development of iPSC derived Cell therapies are exceeding ¥100B<sup>\*</sup>.

\* Financial data is as of Dec 2021.

- Fate Therapeutics (Developing iPSC derived CAR-T/NK products, Market cap: >¥600B)
- Sana Biotechnology (Successful IPO in Feb 2021. Market cap: >¥300B)
- BlueRock Therapeutics (Wholly acquired by Bayer in 2019. Implied total company value: approx. ¥100B)
- Century Therapeutics (Successful IPO in Jun 2021. Market cap: approx. ¥100B)



Establish the Recurring-type of business which enable to achieve sustainable growth by the combination of "Grant of IP license for iPSCs related" and "Process development & manufacturing service utilizing GMP facility (i-FACT)".





#### Contract service and Capital gain from investees



• FCDI invested in Century Therapeutics (found in 2018) to have a contract service for **development and Manufacturing of iPSC-used next-generation immune oncology treatments**.

\* Gain on equiry securities resulting from IPO was posted in June 2021.



• FCDI executed a manufacturing services agreement with Cynata Therapeutics in December 2021 for the manufacture and supply of Cynata's mesenchymal stem cell ("MSC") products treating GvHD.

\* GvHD (Graft versus Host Disease) A general term for a symptom that occurs as a result of the white blood cells present in the organ of a donor recognizing the recipient's body cells as oreign and attacking them.

## Strategic alliance for the development of a in-house pipeline

# 

- A strategic R&D alliance agreement was reached with BlueRock Therapeutics LP regarding iPS cell therapies for ocular diseases.
- An upfront payment of US\$30 million has been received and up to US\$40 million funding for R&D and part of clinical manufacturing is planned to be received for the development of the three retinal disease therapy programs.
- FCDI will also receive milestone payments as development and sales progress, and commercial sales royalty (around 10%).

**GMP Facility "i-FACT"** 



Innovation Facility for Advanced Cell Therapy







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## **Strengthen Drug Discovery & Manufacturing Support**

→ Provide pharma companies and academia with "the solution in the combination of cells, culture media, and reagents" leveraging the expansion of the utilization for the drug discovery screening and pharmacological test by human iPSCs.

## Dramatic growth of cell culture media business

- $\rightarrow$  Establish the global production structure in Japan, US and Europe though continuous Capex.
- → Aim to become a market leader of serum-free culture media for bioproduction (BP) holding 30% share in 2030 by developing customized culture media responding to customers' various needs.

## Expand Cell therapy PD & Mfg Service Business

→ Establish the Recurring-type of business which enable to achieve sustainable growth by "Grant of IP license for iPSCs related" and "Process development & manufacturing service utilizing GMP facility(i-FACT)".

## **Environmental awareness**

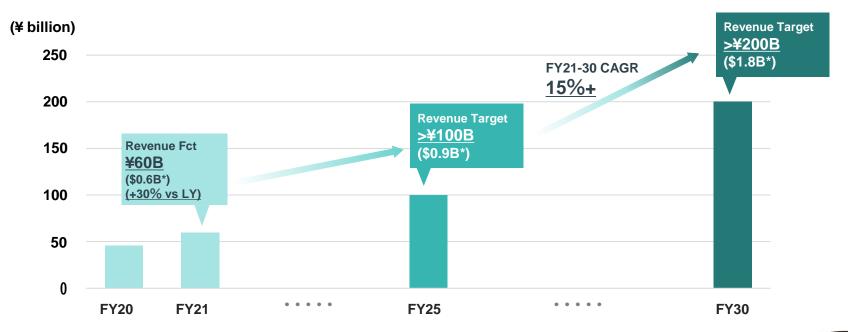
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 $\rightarrow$  Localization of production with 3 sites (U.S., Japan, and Europe) and use of renewable energy to reduce the environmental burden.





#### Aim to reach >¥100B sales in FY25 and >¥200B in FY30 (CAGR 15%+) with BP media as the driving force.



\*109 yen / USD



# FUJ:FILM Value from Innovation

#### Life Science Biz Div Value from Innovation

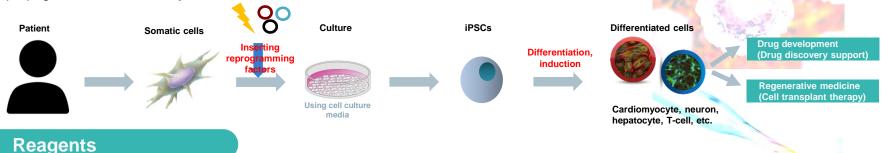
## **Cell Culture Media**

- · Liquid or solid materials that contain nutrients, to be used for culturing microorganisms and cells.
- Culture media consist of basal media (including amino acids, sugars, lipids, vitamins, and salts) and additives (serum or serum-equivalents, some growth factors, etc.).

### **iPS Cells**

(iPS=induced pluripotent stem)

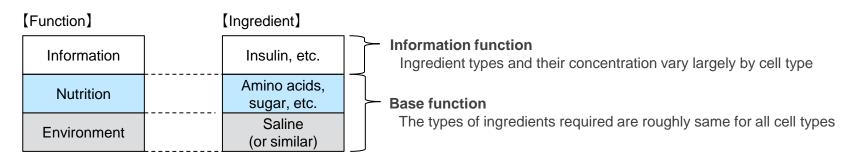
 iPSC stands for induced pluripotent stem cell, produced by introducing a small number of genes known as reprogramming factors to human skin tissues and blood-derived somatic cells to give the ability to differentiate into various tissues and organ cells and the ability to propagate almost indefinitely.



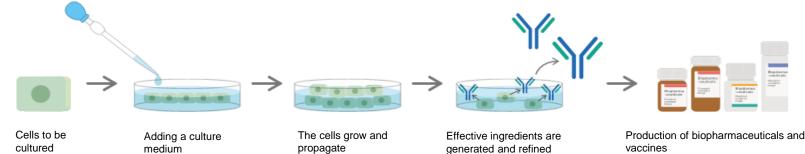
- Reagent refers to a chemical used for testing and research and defined as a "chemical substance used for the detection or qualification of a substance by a chemical process, or for the experimental synthesis of a substance, or for the measurement of the physical characteristics of a substance" (Act on the Evaluation of Chemical Substances and Regulation of Their Manufacturers).
- There are reagents for biochemical research, reagents for genetic engineering research and life-science reagents such as those for immune research.

What are Cell culture media?

Cell culture media are important materials, essential for facilitating cell growth and production of end objects generated from cells. It has the function of providing "environment, nutrients and information" to cells. Just as people have personal preferences, cells and cell products have individual preference in optimum composition of culture media.



Cell culture process





Life Science Biz Div

# NEVER STOP

Achieving Continual Growth