



REALIZING THE POTENTIAL OF PLASMA-DERIVED THERAPIES

21st November 2019

Julie Kim

President, Plasma-Derived Therapies Business Unit



IMPORTANT NOTICE



For the purposes of this notice, "presentation" means this document, any oral presentation, any question and answer session and any written or oral material discussed or distributed by Takeda Pharmaceutical Company Limited ("**Takeda**") regarding this presentation. This presentation (including any oral briefing and any question-and-answer in connection with it) is not intended to, and does not constitute, represent or form part of any offer, invitation or solicitation of any offer to purchase, otherwise acquire, subscribe for, exchange, sell or otherwise dispose of, any securities or the solicitation of any vote or approval in any jurisdiction. No shares or other securities are being offered to the public by means of this presentation. No offering of securities shall be made in the United States except pursuant to registration under the U.S. Securities Act of 1933, as amended, or an exemption therefrom. This presentation is being given (together with any further information which may be provided to the recipient) on the condition that it is for use by the recipient for information purposes only (and not for the evaluation of any investment, acquisition, disposal or any other transaction). Any failure to comply with these restrictions may constitute a violation of applicable securities laws.

The companies in which Takeda directly and indirectly owns investments are separate entities. In this presentation, "Takeda" is sometimes used for convenience where references are made to Takeda and its subsidiaries in general. Likewise, the words "we", "us" and "our" are also used to refer to subsidiaries in general or to those who work for them. These expressions are also used where no useful purpose is served by identifying the particular company or companies.

Forward-Looking Statements

This presentation and any materials distributed in connection with this presentation may contain forward-looking statements, beliefs or opinions regarding Takeda's future business, future position and results of operations, including estimates, forecasts, targets and plans for Takeda. Without limitation, forward-looking statements often include words such as "targets", "plans", "believes", "hopes", "continues", "expects", "aims", "intends", "ensures", "will", "may", "should", "would", "could" "anticipates", "estimates", "projects" or similar expressions or the negative thereof. Forward-looking statements in this document are based on Takeda's estimates and assumptions only as of the date hereof. Such forward-looking statements do not represent any guarantee by Takeda or its management of future performance and involve known and unknown risks, uncertainties and other factors, including but not limited to: the economic circumstances surrounding Takeda's global business, including general economic conditions in Japan and the United States; competitive pressures and developments; changes to applicable laws and regulations; the success of or failure of product development programs; decisions of regulatory authorities and the timing thereof; fluctuations in interest and currency exchange rates; claims or concerns regarding the safety or efficacy of marketed products or product candidates; the timing and impact of post-merger integration efforts with acquired companies; and the ability to divest assets that are not core to Takeda's operations and the timing of any such divestment(s), any of which may cause Takeda's actual results, performance, achievements or financial position to be materially different from any future results, performance, achievements or financial position expressed or implied by such forward-looking statements. For more information on these and other factors which may affect Takeda's results, performance, achievements, or financial position, see "Item 3. Key Information—D. Risk Factors" in Takeda's most recent Annual Report on Form 20-F and Takeda's other reports filed with the U.S. Securities and Exchange Commission, available on Takeda's website at: <https://www.takeda.com/investors/reports/sec-filings/> or at www.sec.gov. Future results, performance, achievements or financial position of Takeda could differ materially from those expressed in or implied by the forward-looking statements. Persons receiving this presentation should not rely unduly on any forward-looking statements. Takeda undertakes no obligation to update any of the forward-looking statements contained in this presentation or any other forward-looking statements it may make, except as required by law or stock exchange rule. Past performance is not an indicator of future results and the results of Takeda in this presentation may not be indicative of, and are not an estimate, forecast or projection of Takeda's future results.

Medical information

This presentation contains information about products that may not be available in all countries, or may be available under different trademarks, for different indications, in different dosages, or in different strengths. Nothing contained herein should be considered a solicitation, promotion or advertisement for any prescription drugs including the ones under development.

Financial information

Takeda's financial statements are prepared in accordance with International Financial Reporting Standards ("IFRS").

The revenue of Shire plc ("Shire"), which were presently, presented in accordance with accounting principles generally accepted in the United States ("U.S. GAAP"), have been conformed to IFRS, without material difference.

The Shire acquisition closed on January 8, 2019, and our consolidated results for the fiscal year ended March 31, 2019 include Shire's results from January 8, 2019 to March 31, 2019. References to "Legacy Takeda" businesses are to our businesses held prior to our acquisition of Shire. References to "Legacy Shire" businesses are to those businesses acquired through the Shire acquisition.

This presentation includes certain pro forma information giving effect to the Shire acquisition as if it had occurred on April 1, 2018. This pro forma information has not been prepared in accordance with Article 11 of Regulation S-X. This pro forma information is presented for illustrative purposes and is based on certain assumptions and judgments based on information available to us as of the date hereof, which may not necessarily have been applicable if the Shire acquisition had actually happened as of April 1, 2018. Moreover, this pro forma information gives effect to certain transactions and other events which are not directly attributable to the Shire acquisition and/or which happened subsequently to the Shire acquisition, such as divestitures and the effects of the purchase price allocation for the Shire acquisition, and therefore may not accurately reflect the effect on our financial condition and results of operations if the Shire acquisition had actually been completed on April 1, 2018. Therefore, undue reliance should not be placed on the pro forma information included herein.


PLASMA PRESENTS A UNIQUE OPPORTUNITY




Plasma is a durable business with compelling growth opportunity...

- 1 Products have lifecycle spanning decades
- 2 Indication expansion continues
- 3 Not subject to patent cliffs
- 4 Probability of success for R&D is generally high
- 5 Market demand continues to grow steadily

...AND HAS DISTINCT ASPECTS



Plasma is collected from human donations - scarce supply



It can take more than 7 months to produce plasma-derived therapies



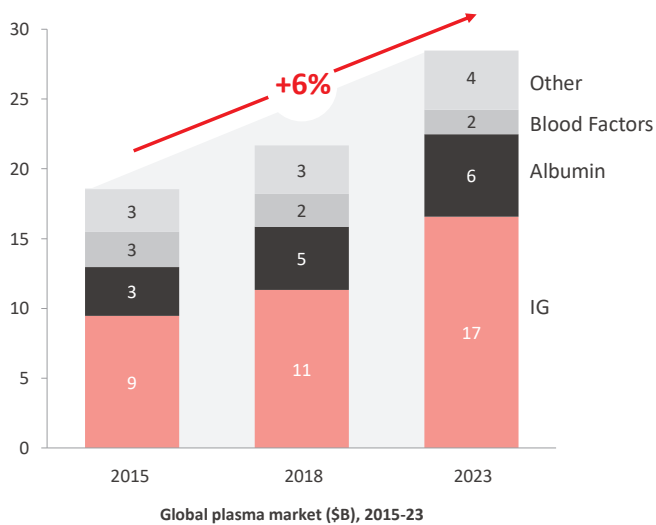
Capital-intensive manufacturing process

2





DEMAND FOR KEY PLASMA-DERIVED THERAPIES HAS BEEN CONTINUOUSLY INCREASING AND EXPECTED TO GROW



Worldwide demand for plasma-derived therapies is expected to increase each year



This trend is primarily driven by:

-  Greater awareness and increasing rates of diagnosis
-  Growing access in emerging markets
-  New indications in both immune deficiencies and immune-mediated diseases
-  Innovation in formulations and delivery systems

Source: Berman. Plasma Fractionation: The Challenge of Keeping Pace with Global IG Demand / MRB, Evaluate Pharma, PDT Analysis / Chapel H, et al. Front Immunol 2014 Dec 15;5:627. / News release: Shire Launches Paediatric Indication for Immunodeficiency Treatment HyQvia in Europe. 21 July 2016. / News release: Shire Announces FDA Approval for Label Expansion of CINRYZE® for Prevention of Attacks in Pediatric HAE / Jones, et al. Frontiers in Immunology 2018;9:1308.

TAKEDA IS NOW ORGANIZED – AND UNIQUELY POSITIONED - TO REALIZE THE FULL POTENTIAL OF PLASMA-DERIVED THERAPIES



>20

PLASMA-DERIVED THERAPIES

RARE DISEASE LEADER

Deep understanding of patient's needs

GLOBAL PHARMA SCALE & EXPERTISE

Capabilities in digital technology, data analytics, patient insights

75+ YEAR

pioneer legacy in plasma

PLASMA-DERIVED THERAPIES DEDICATED BUSINESS UNIT

Top 3 plasma company, investing to grow



PLASMA-FOCUSED R&D Team

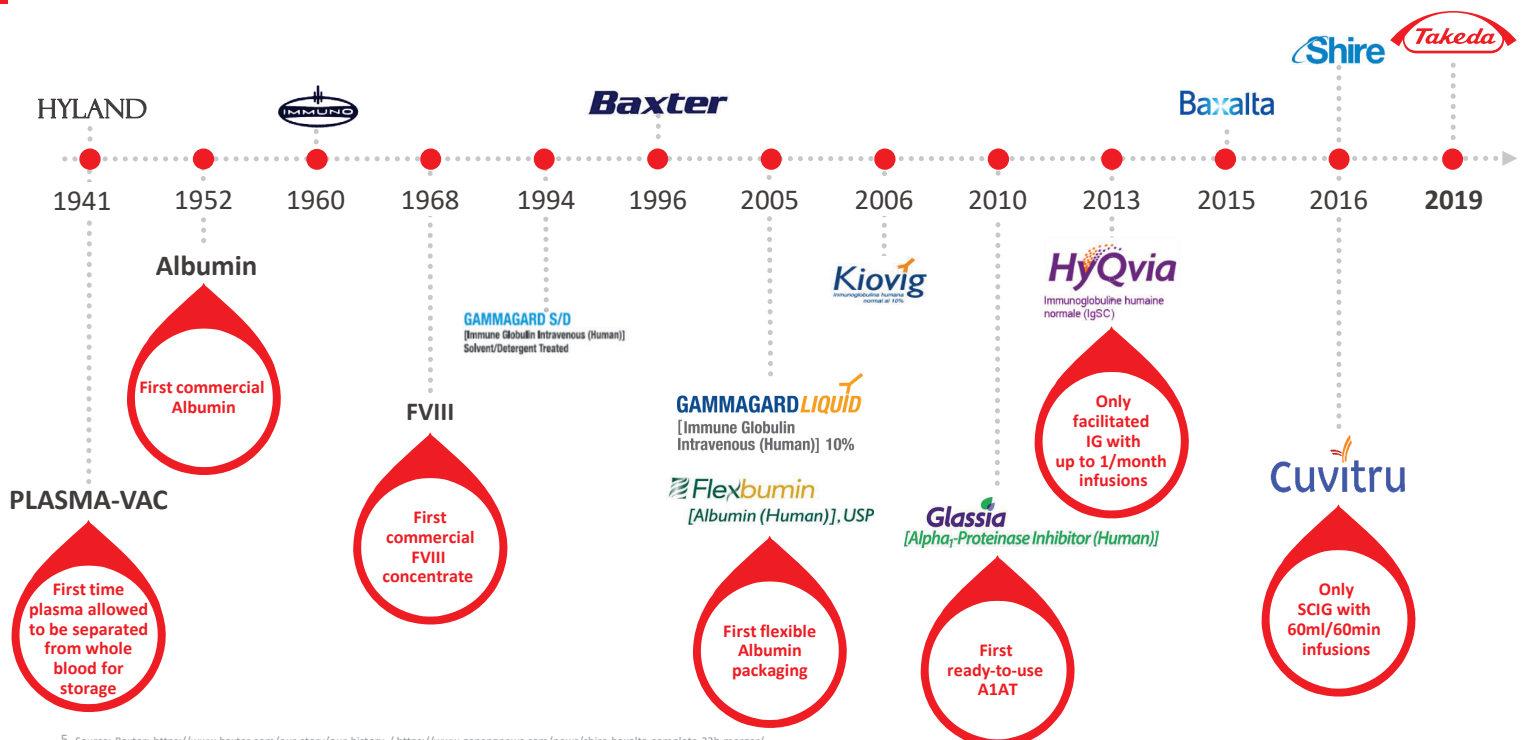
8 MANUFACTURING SITES

140+ PLASMA COLLECTION CENTERS

13,000 EMPLOYEES worldwide, focused on plasma business

4 Source: Evaluate Pharma, PDT Analysis. / Takeda internal data

WE ARE BUILDING ON A LONG AND SUCCESSFUL HISTORY OF BRINGING INNOVATIVE THERAPIES TO PATIENTS



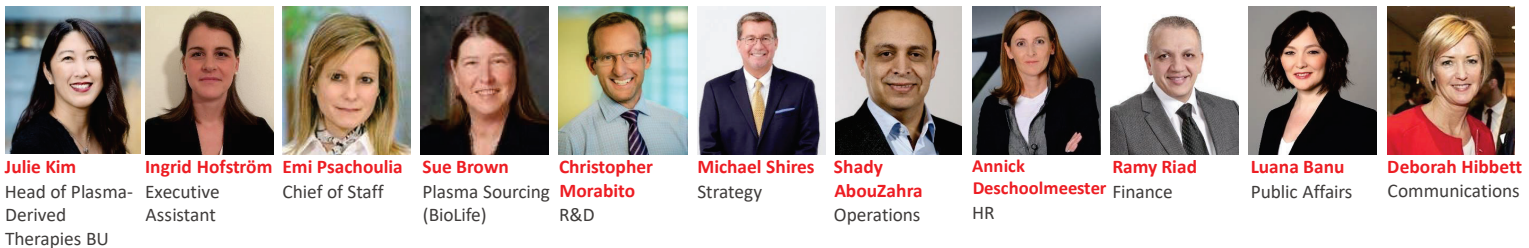
5 Source: Baxter: <https://www.baxter.com/our-story/our-history> / <https://www.genengnews.com/news/shire-baxalta-complete-32b-merger/>

WE'VE ESTABLISHED A DEDICATED BUSINESS UNIT TO STEER OUR PATH, BRING FOCUS AND HARNESS OUR END-TO-END PLASMA CAPABILITIES



6

OUR PDT BU LEADERSHIP TEAM DRAWS ON, AND BRINGS TOGETHER, TAKEDA'S EXTENSIVE PLASMA EXPERIENCE AND BROADER EXPERTISE



Julie Kim Head of Plasma-Derived Therapies BU
Ingrid Hofström Executive Assistant
Emi Psachoulia Chief of Staff
Sue Brown Plasma Sourcing (BioLife)
Christopher Morabito R&D
Michael Shires Strategy
Shady AbouZahra Operations
Annick Deschoolmeester HR
Ramy Riad Finance
Luana Banu Public Affairs
Deborah Hibbett Communications



Adrian Murphy Manufacturing
Barbara Glantschnig Quality
Thomas Kreil Pathogen Safety
Kasha Witkos Commercial
Paula Leca Legal
Gabriele Ricci IT
Linda Peralta Ethics & Compliance
Charlie Alexander Business Development

140 +
combined years of plasma experience

19
team members

11
nationalities

♀ 60% Female
♂ 40% Male

7

Our Ambition

Build a respected, sustainable plasma business that reimagines the industry to best serve patients worldwide



8

We have a singular, dedicated plasma focus and strategy

- Responsibility for end-to-end plasma business
- Dedicated R&D organization and budget

We also benefit from the support of a global, values-based biopharmaceutical company

- Long-term view with commitment to invest as plasma is a key growth driver for Takeda
- Access to Takeda's broader resources, capabilities and expertise, particularly R&D and manufacturing

OUR STRATEGY AND TARGETED INVESTMENTS EXTEND ACROSS THE ENTIRE VALUE CHAIN

PLASMA SOURCING

MANUFACTURING

COMMERCIALIZATION

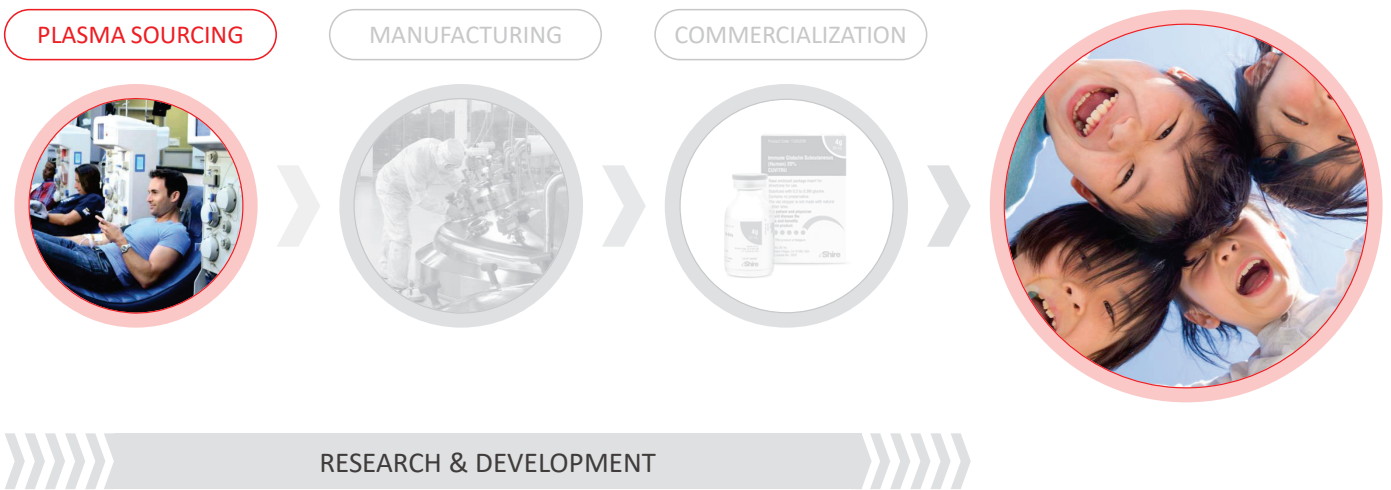


RESEARCH & DEVELOPMENT



9

VIDEO OF PLASMA CENTER AND MANUFACTURING FACILITY



WE ARE ACCELERATING THE RATE OF PLASMA COLLECTION AND INCREMENTALLY INCREASING OVERALL VOLUME



We are building momentum....



- Increased plasma volumes by approximately 20% in 2018
- Expanded European presence from 7 to 30 collection centers within past 12 months
- Completed 5 acquisitions in the past 12 months in US, Austria, Hungary and Czechia
- Plan on opening a total of 19 additional new collection centers in fiscal year 2019
- Leveraging third party supply through long-term contracts
- Participating in contract agreements with governments

We will continue to focus on operational excellence



- Open collection sites faster
- Increase speed to peak collection volumes
- Create efficiency via new models and approaches

We are accelerating growth with the goal of increasing plasma supply by

>65%

over the next 5 years

12 Source: Takeda internal data.

WE ARE FURTHER ENHANCING AND DIGITALIZING FACILITIES AND SERVICES TO MEET GROWING NEEDS FOR THE FUTURE



Attracting new donors in the community

- Reaching new donors
- Increasing community engagement

Improving the donor experience and improving cost-per-liter through omnichannel engagement



Mobile App



Website



Scheduling



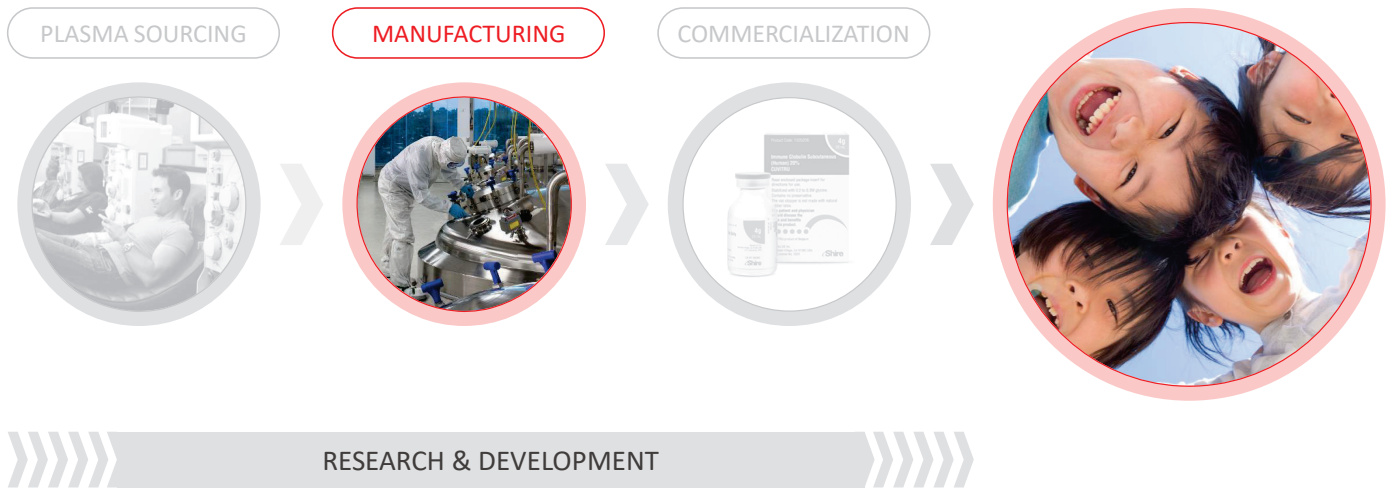
Payment



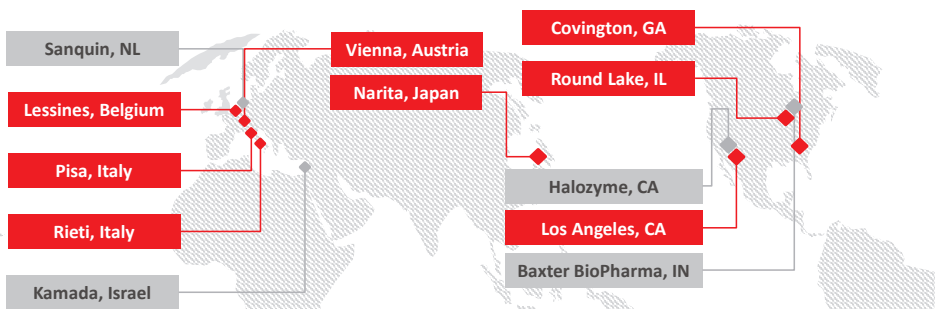
Information

Donor





WE HAVE A WORLD-LEADING PLASMA DERIVED THERAPIES MANUFACTURING NETWORK IN WHICH WE CONTINUE TO SIGNIFICANTLY INVEST



8 STRATEGIC LOCATIONS

plus four strategic partners, allowing independent yet inter-related manufacturing operations

INNOVATION MINDSET

digitalization and constant drive for excellence to accelerate supply to patients

CONTINUED CAPACITY EXPANSION

to increase production of our portfolio to meet market growth while driving efficiencies

CONTINUALLY INVESTING

in state-of-the-art facilities that meet the highest quality standards



Takeda Mfg.

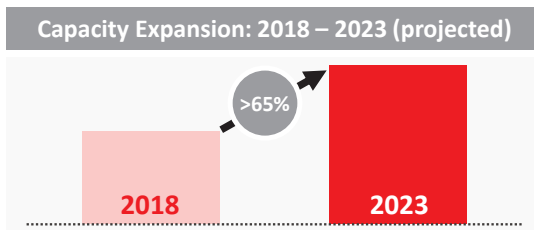
External Mfg.

WE'RE INCREASING PRODUCTION CAPACITY BY ACCELERATING INVESTMENT, WHILE FURTHER ENHANCING OUR QUALITY STANDARDS



Investing in manufacturing capacity

- Continually investing in technologies and processes to **maximize yield**
 - Higher yield, lower cost fractionation techniques
 - Analytics, automation and digitization to optimize network
- **Optimizing plasma efficiency** through the value chain
- **Downstream optimization** within broader Takeda manufacturing network



We plan to increase our manufacturing capacity within our existing network by

>65%

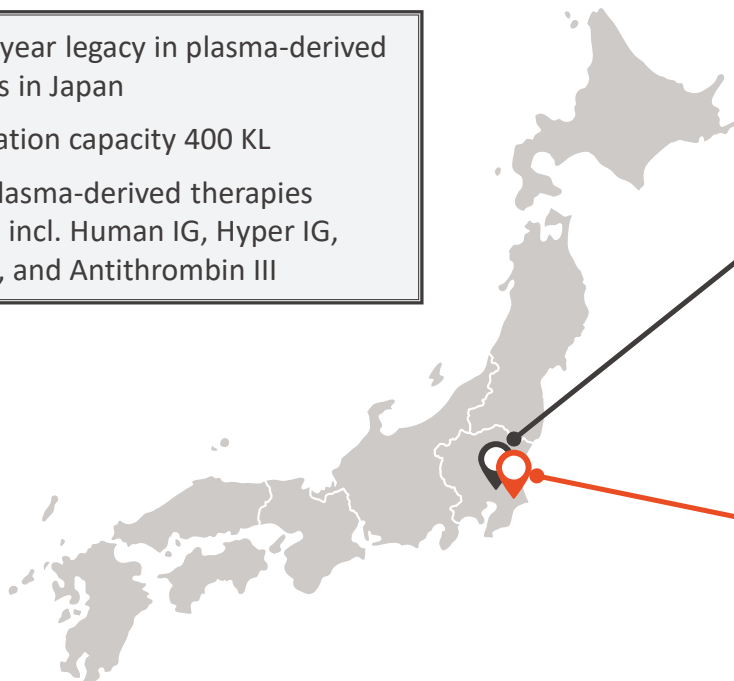
over the next 5 years

16

IN JAPAN TAKEDA HAS DOMESTIC MANUFACTURING CAPABILITIES THROUGH OUR SUBSIDIARY NIHON PHARMA



- Over 60-year legacy in plasma-derived therapies in Japan
- Fractionation capacity 400 KL
- Strong plasma-derived therapies portfolio incl. Human IG, Hyper IG, Albumin, and Antithrombin III



17

TAKEDA HAS WORLD-CLASS SAFETY CAPABILITIES AND AN UNSURPASSED REPUTATION IN BOTH PLASMA DONATION AND PATHOGEN SAFETY



Donation safety standards

Strict donation criteria and screening at each visit

Donation frequency management system

Strong inspection record

Plasma screening, inventory hold and look back procedure

Every plasma donation screened for HIV, hepatitis A, B & C, parvo B19

Pathogen safety standards

BioSafety Level 3+ Lab

Purpose-built, state-of-the-art biocontainment laboratory

Process sciences

Qualified models of all bioprocessing steps

Virology

Classical & molecular virology expertise and capability

Publication / presentation

Strong track record

Dedicated virology expertise and capabilities



40+ highly trained staff



>50% with specialized education



>200 years post-graduate experience



PLASMA SOURCING

MANUFACTURING

COMMERCIALIZATION



RESEARCH & DEVELOPMENT

OUR BROAD AND DIFFERENTIATED PORTFOLIO OF PLASMA-DERIVED THERAPIES TREATS RARE AND COMPLEX DISEASES WORLDWIDE



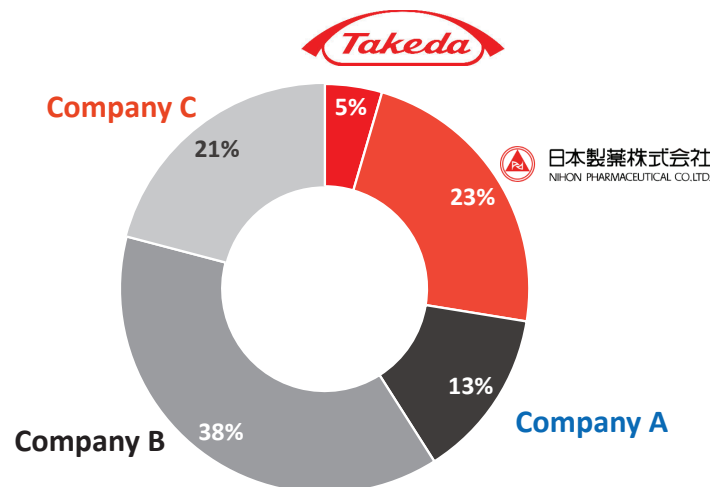
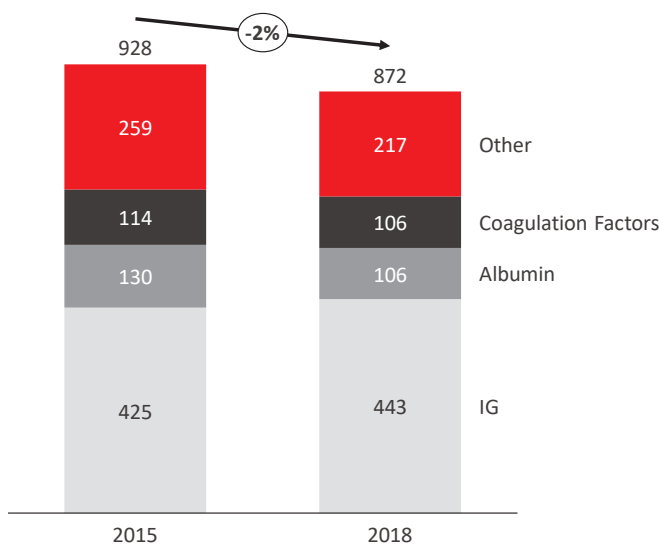
20 For illustrative purposes only, geographies and products do not correspond

IN JAPAN, IG GROWTH REMAINS STRONG AND THERE IS OPPORTUNITY TO ENHANCE STANDARD OF CARE



Japan Plasma Market (\$M), 2015-18

Japan PDT Revenue Market Share



Pd: plasma derived, IG: Immunoglobulin

NOTE: The category of product type is based on Takeda internal standards. Converted at April 2018-March 2019 average exchange rate of 111 JPY/USD
 Copyright © 2019 IQVIA. (Calculated based on JPM Jan - Dec 2013 and Jan - Dec 2018) Reprinted with permission.

Copyright © 2019 IQVIA. Calculated based on JPM Jan - Dec 2013 and Jan - Dec 2018. Reprinted with permission.

OUR GOAL IS TO CONTINUE TO BRING PERSONALIZED, INNOVATIVE, LIFELONG CARE THROUGHOUT THE PATIENT JOURNEY



Diagnosis

- Partnership with large hospital systems in the US to leverage electronic medical records
- Co-chairing the Global Commission to End the Diagnostic Odyssey for Children with Rare Disease
- Awareness campaigns
- Diagnostic test kits



Access

- Sustainable pricing
- Dedicated access support
- Patient assistance programs
- Broad portfolio of products

Personalized Care & Support

- Enhanced patient services
- Nurse training to support new patients
- Devices and delivery systems

22

WE ANTICIPATE SIGNIFICANT GROWTH OPPORTUNITIES ACROSS OUR PORTFOLIO



		Example Global Takeda products	Japan Takeda products	Global Takeda revenue (OY, 2018)	Global plasma market size (OY, 2018)
Last Liter	Immunoglobulin	GAMMAGARD <i>Liaqto</i> / <i>Kiovig</i> HyQvia Cuvitru	kenketu glovenin [®] GAMMAGARD S/D	~2,870	~12,500
	Albumin	Flexbumin HUMAN ALBUMIN	KENKETU ALBUMIN KENKETU ALBUMINATE [®]	~580	~5,000
First Liter	Hemophilia products	HEMOFIL M IMMUNINE [®]	FEIBA [®] IMMUNATE	~890	~2,800
	Other products	Aralast NP Glassia Prothromplex NF 600 Ceprotin Antithrombin III CINRYZE [®]	KENKETU NONTHRON [®] Antithrombin III	~660	~3,700
Total				~5,000*	~24,000

*2018 revenue is a pro-forma which adds Legacy Shire's 9 month (April - December 2018) revenue previously reported under US GAAP and conformed to IFRS without material differences and converted to JPY using FY2018 actual rate for the period. 2018 revenue also includes product sales of Nihon Pharmaceutical products, Takeda's consolidated subsidiary.

23

Source: MRB; EvaluatePharma; Takeda internal data

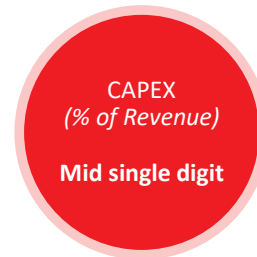
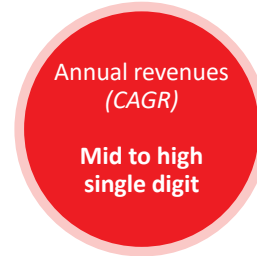
AND WE ARE EMBARKING ON A TRAJECTORY TO IMPROVE OVERALL PLASMA-DERIVED THERAPIES BUSINESS PERFORMANCE



Key Growth & Margin Drivers for PDT

- Focused **sustainable, value-based commercial strategies**, including tenders
- **Process efficiencies** across the network
- **Capacity increase** across collections and manufacturing
- **R&D investments** across portfolio

Key Financial Aspiration for PDT*



* The "Key Financial Aspirations" listed above represent Takeda's goals in the long-term for the PDT business as of the date hereof and are based on certain assumptions. Actual Amounts/results may differ materially and are subject to a number of risks and uncertainties. See "Note Regarding Forward Looking Statements" on Page 1 of this presentation.

KEY TAKEAWAYS



1

At Takeda, plasma is a **long-term strategic focus**, led by a **dedicated business unit investing to grow** across the value chain and leveraging Takeda capabilities

2

Our goal is to **accelerate growth in capacity by >65%** over the next 5 years to bring additional and improved therapies to more people around the world

3

Our **broad and differentiated portfolio** brings **personalized, innovative, lifelong care** and underlines our credentials for **reimagining the industry**



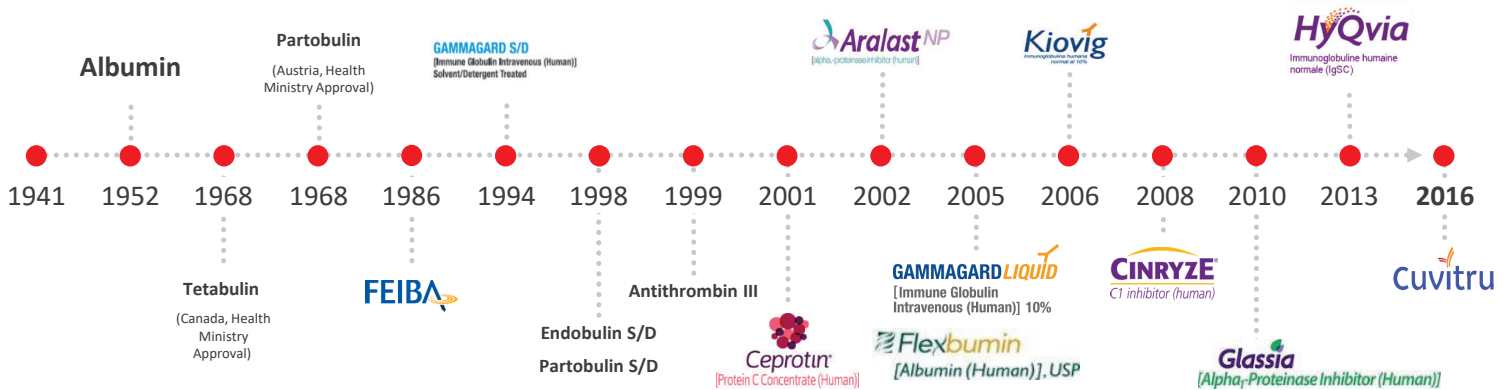
A NEW DEDICATED FOCUS ON INNOVATIVE, SUSTAINABLE SOLUTIONS FOR PLASMA-DERIVED THERAPIES



Christopher Morabito, M.D.
Head of R&D, Plasma-Derived Therapies

Better Health, Brighter Future

PDT R&D'S CREDENTIALS AND INFRASTRUCTURE ARE WELL-ESTABLISHED

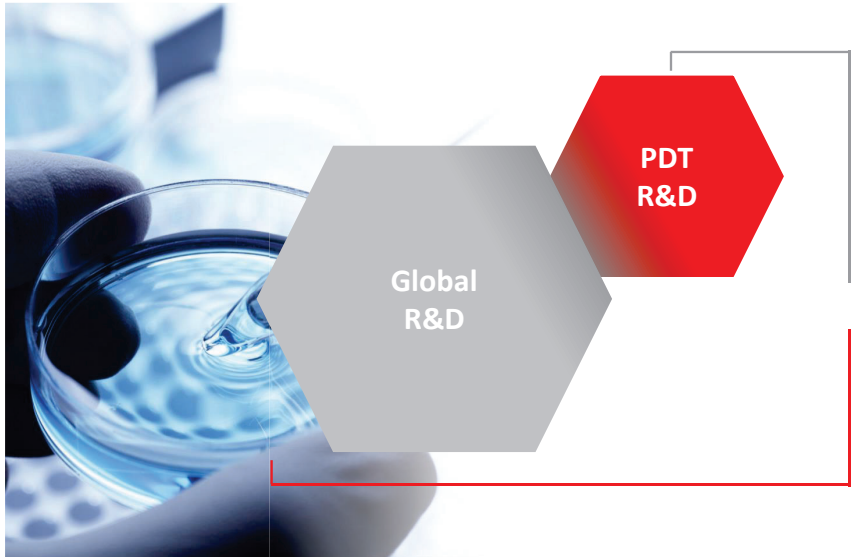


Pathogen Safety
Global Center of Excellence for Pathogen Safety

Pharmaceutical Science
Strong team connected across the value chain

Pilot Labs
Within Vienna, Los Angeles, Georgia and Lessines sites

OUR INDEPENDENCE BRINGS FOCUS ON PLASMA AND IS BOLSTERED BY ACCESS TO BROADER R&D CAPABILITIES AND RESOURCES



- Focused entirely on plasma-derived therapies
- Lean and agile team
- Based in Cambridge, MA and Vienna, Austria
- Separate R&D prioritization
- Dedicated budget

- Common Takeda values, patient-focused vision
- Common governance
- Shared resources (e.g. Medical Affairs, Safety, Quality)

These links strengthen Takeda R&D's modality mix, now the broadest among the Top 10 global biopharmaceutical companies

THE PDT R&D LEADERSHIP TEAM IS WELL-INTEGRATED AND BRINGS DEEP AND DIVERSE FUNCTIONAL EXPERTISE



Christopher Morabito MD
R&D Head
Boston, MA



Catherine Parham MD
Program Leadership
Boston, MA



Rory Bukofzer
Program Leadership
Boston, MA



Leman Yel MD
Clinical Medicine
Boston, MA



Chris Tremblay
R&D Operations
Boston, MA



Bagirath Gangadharan PhD
Translational Research
Vienna, Austria



Andreas Liebming PhD
Pharmaceutical Sciences
& Devices
Vienna, Austria/Boston, MA



Sascha Haverfield DPhil
Regulatory Affairs &
Development Operations
Boston, MA



Geoffrey Pot PhD
Global Manufacturing
External Supply & Plasma
Innovation
Lessines, Belgium



Gabriele Ricci
Digital Technologies
Boston, MA



William Standaert
Legal
Zurich, Switzerland



Cara Laurello
Ethics and Compliance
Boston, MA



Ambreen Landa
Human Resources
Boston, MA

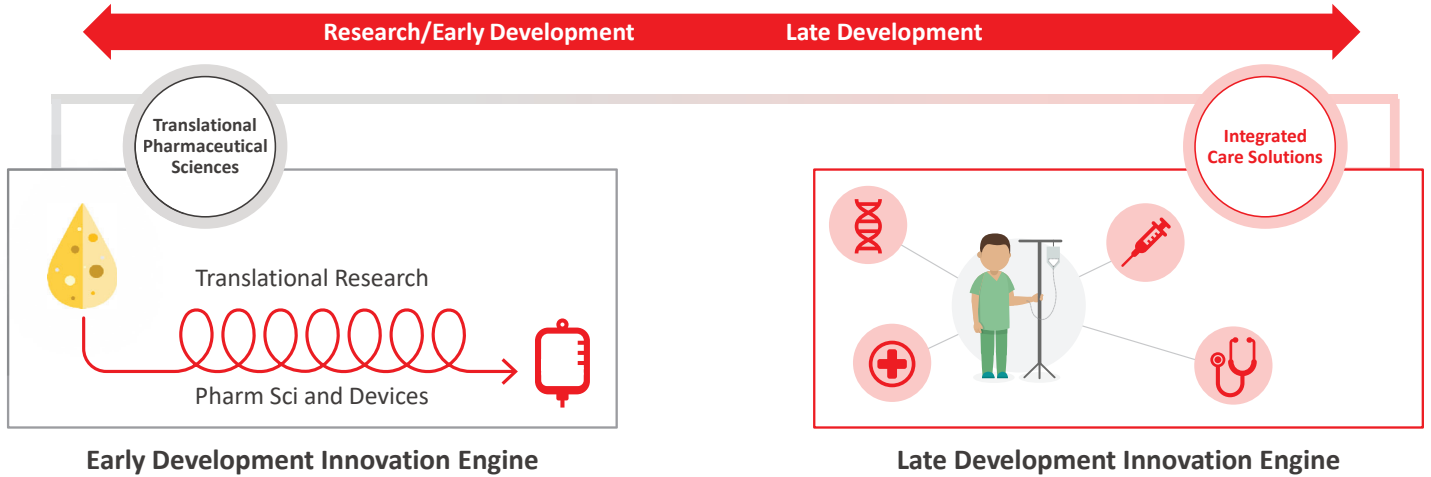


Pritesh Patel
Finance
Boston, MA



Julia Ellwanger
Communications
Bannockburn, IL

WE ARE DRIVING A CULTURE OF INNOVATION THROUGH TWO R&D ENGINES



- Generate new and improved therapeutics by:**
- Investigational new drug candidates
 - Mechanisms of action
 - Responder populations
 - New process development

- Improve health outcomes by:**
- Diagnostic efficiencies
 - Expanded data and devices to support effectiveness
 - Point of Care services and drug delivery services
 - Data-driven guidelines for acute and chronic management

PDT R&D Strategy

Maximize the therapeutic value of plasma-derived therapies for patients with rare and complex diseases through innovation across the product life cycle



Realize full potential of in-line First and Last Lifer products

- Expanded indications and benefit-risk datasets
- Device-driven solutions for diagnosis, management, and long-term follow-up
- Global expansion
- New formulations



Optimize efficiencies of plasma-derived therapy production

- Pharmaceutical science support for manufacturing



Identify and develop new plasma-derived therapies

- New targeted therapies for diverse therapeutic areas

WE ARE PRIORITIZING NEAR-TERM LATE DEVELOPMENT...



	RESEARCH / NON-CLINICAL DEVELOPMENT	LATE DEVELOPMENT
IMMUNOLOGY	<p>CUVITRU Wearable Device</p>	<p>HYQVIA <i>Halozyme</i> US - Pediatric PID</p> <p>HYQVIA <i>Halozyme</i> Chronic inflammatory demyelinating polyneuropathy (CIDP)</p> <p>HYQVIA Geographic expansion</p> <p>CUVITRU Geographic expansion</p>
		<p>HYQVIA <i>Halozyme</i> EU - Pediatric PID</p> <p>HYQVIA - HyHub <i>Flextronics</i> Delivery Device</p> <p>CINRYZE Geographic expansion</p> <p>GLASSIA <i>Kamada</i> Immunogenicity/ bronchioalveolar lavage</p>
HEMATOLOGY		<p>FEIBA Volume reduction</p>

32

... WHILE ENABLING DISCOVERY OF NEXT GENERATION THERAPEUTICS



	RESEARCH / NON-CLINICAL DEVELOPMENT	LATE DEVELOPMENT
IMMUNOLOGY	<p>CUVITRU Wearable Device</p> <p>TAK 880 Low IgA-IgG (IV) Primary Immunodeficiency</p> <p>Hyper-Immune IG Infectious disease</p> <p>CINRYZE Ex-HAE indications TBD</p>	<p>TAK 881 Facilitated 20% SC IgG <i>Halozyme</i> Primary Immunodeficiency (PID)</p> <p>Alpha-1 Antitrypsin (A1AT) Next generation formulations</p> <p>HYQVIA <i>Halozyme</i> US - Pediatric PID</p> <p>HYQVIA <i>Halozyme</i> Chronic inflammatory demyelinating polyneuropathy (CIDP)</p> <p>HYQVIA Geographic expansion</p> <p>CUVITRU Geographic expansion</p> <p>GLASSIA <i>Kamada</i> A1ATD-empysema*</p>
		<p>HYQVIA <i>Halozyme</i> EU - Pediatric PID</p> <p>HYQVIA - HyHub <i>Flextronics</i> Delivery Device</p> <p>CINRYZE Geographic expansion</p> <p>GLASSIA <i>Kamada</i> Immunogenicity/ bronchioalveolar lavage</p> <p>CUVITRU Japan - PID (FPI Q4 2019)</p>
HEMATOLOGY	<p>PROTHROMPLEX TOTAL Device and formulation</p> <p>Butyryl Cholinesterase Organophosphate poisoning</p>	<p>PROTHROMPLEX TOTAL US - Drug-induced bleeding**</p> <p>CEPROTIN Geographic expansion</p> <p>FEIBA Volume reduction</p>

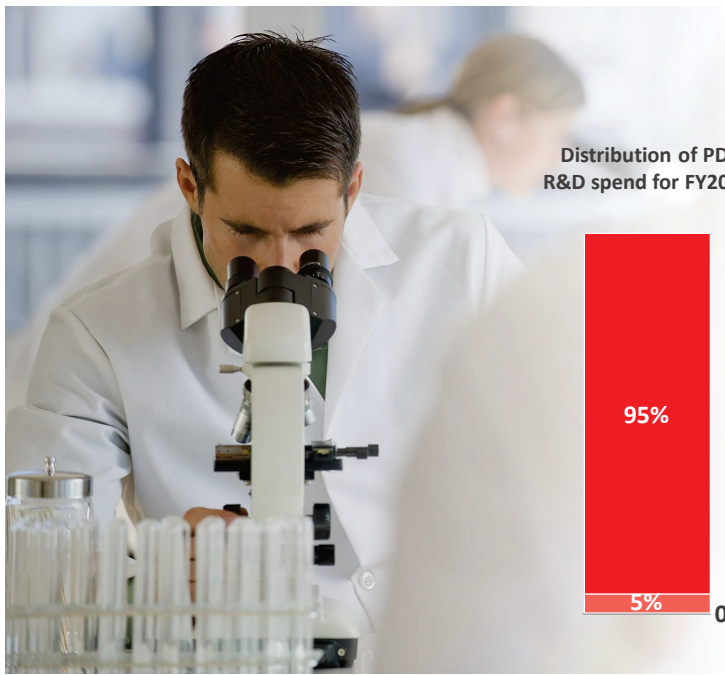
33

*Subject to regulatory approval

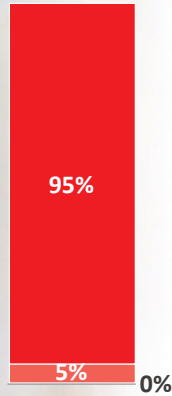
**Pending FDA Pre-IND consultation and future acceptance of an IND

Programs and projects added since Day 1

OVER THE NEXT 3 YEARS, WE PLAN TO ALLOCATE RESOURCES TO RESEARCH AND EARLY DEVELOPMENT



Distribution of PDT R&D spend for FY2019



Estimated % of PDT R&D spend for FY2023



~70% of resources will be allocated to improving in-line products and production efficiencies



Optimizing value of in-line products



Plasma production efficiencies



New plasma-derived therapies



OUR GOAL IS TO REALIZE THE FULL POTENTIAL OF IN-LINE FIRST AND LAST LITER PRODUCTS



Estimated % of PDT R&D spend for FY2023



- Expanded indications and benefit-risk datasets
- Device-driven solutions for diagnosis, management, and long-term follow-up
- Global expansion
- New formulations



Optimizing value of in-line products



Plasma production efficiencies



New plasma-derived therapies



IMMUNOGLOBULINS PROVIDE THE SCAFFOLD FOR PDT INNOVATION



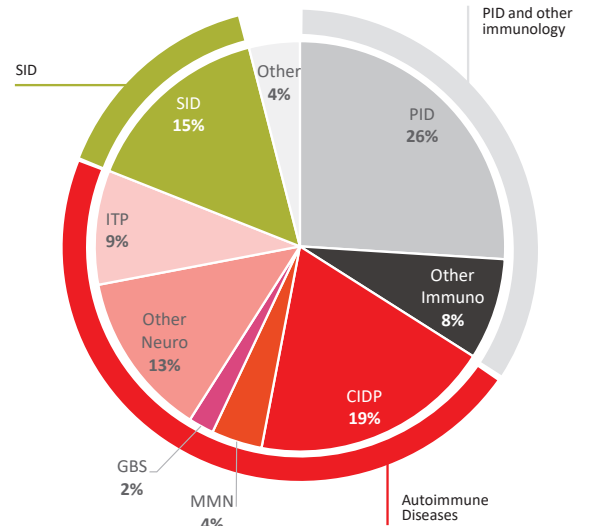
Current State

- Exploring efficacy and safety of HYQVIA in patients with neuro-immune diseases (e.g. CIDP)
- Ongoing delivery device development

Opportunities

- Indications: New neuro-immunology and secondary immunodeficiencies (SID) programs**
- Geographic expansion: CUVITRU-Japan first patient to be enrolled in Q4 FY 2019
- Integrated care solutions:
 - Advance point of care diagnosis of primary immunodeficiency (PID)
 - New delivery and eHealth devices
- Develop f-20% SCIG

US & EU IgG use by indication*



Source: Bain Study (US&EU), Volumes, Estimates based on internal calculations on EU Country Data

*Not all indications are approved for a Takeda product

**Subject to regulatory approval

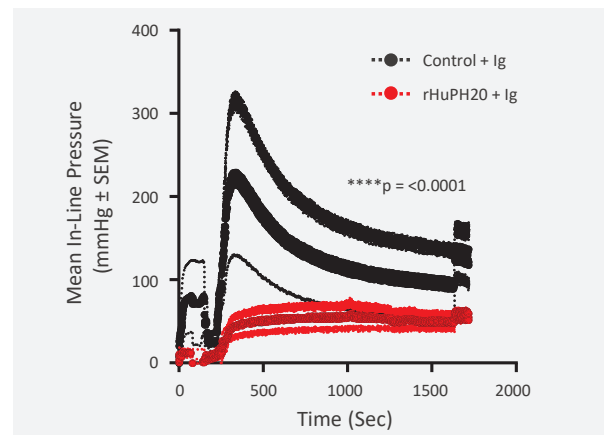
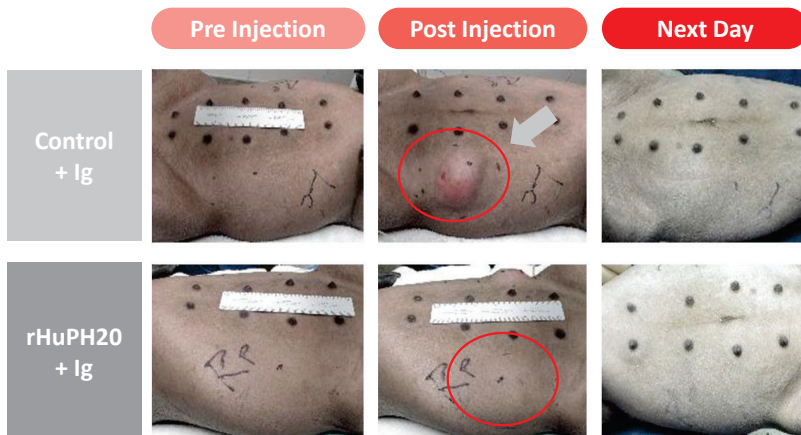
36



FACILITATED 20% SCIG HAS THE POTENTIAL TO PROVIDE FURTHER VALUE TO PATIENTS WHO REQUIRE HIGHER VOLUME ADMINISTRATIONS



Pig model, sequentially administered recombinant human hyaluronidase (rHuPH20) and 20% IgG (CUVITRU)*



**Significantly decreased induration and infusion pressure,
with improved cutaneous blood flow**

* In collaboration with Halozyme

Sequentially administered rHuPH20 and CUVITRU is for investigational use only

37



PROTHROMPLEX TOTAL CAN BE DEVELOPED TO TREAT A VARIETY OF BLEEDING DISORDERS



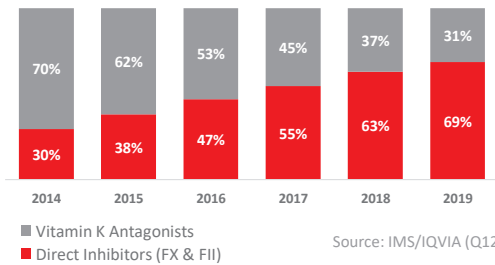
Current State

- Many different mechanisms used for prophylactic and surgical anti-coagulant therapy
- PROTHROMPLEX TOTAL use is limited to Vitamin K antagonists associated bleeding ex-US

Opportunities

- Geographic expansion into the US*
- Broaden indication to include treatment of multiple types of drug-induced bleeding
- Improved use via new formulations and device

Changing Treatment Paradigm
(EU Total Prescriptions)



38 *Pending FDA Pre-IND consultation and future acceptance of an IND; Investigational use, subject to regulatory approval



ARALAST & GLASSIA PROVIDE OPPORTUNITIES TO IMPROVE OUTCOMES IN PATIENTS WITH ALPHA-1 ANTITRYPSIN DEFICIENCY (A1ATD)



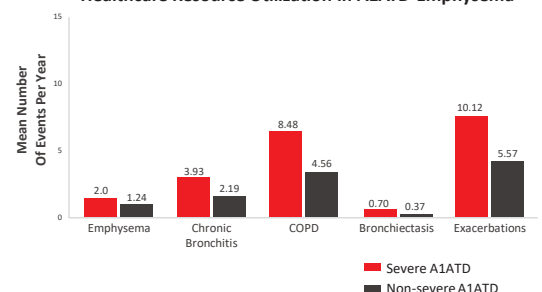
Current State

- Current standard of care does not adequately treat A1ATD

Opportunities

- New clinical study to assess the efficacy of a higher dose of GLASSIA in patient with emphysema related to A1ATD
- Next generation A1AT*: formulation, delivery and management devices
- Explore A1AT as acute phase reactant

Healthcare Resource Utilization in A1ATD-Emphysema



Source: Herrera et al (2019) Chest annual meeting

39 *Investigational use, subject to regulatory approval



INVESTIGATIONAL A1AT-REPLACEMENT FORMULATIONS MAY OFFER ADDITIONAL VALUE TO PATIENTS



Short term

Highly purified post-fractionations
pdA1AT-precursor



Concentration
of A1AT by ultra filtration potentially leading to an **extended $t_{1/2}$**

Formulation Development
Evaluate SC administration

Device Development
Potential to add incremental value for patients

Mid term

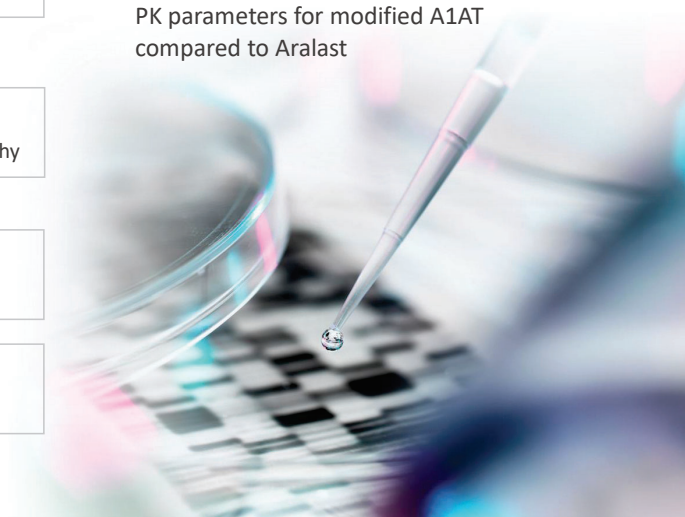
Protein Modification
site-specific modification leading to an **extended $t_{1/2}$**



Purification
by ion-exchange chromatography

In Vivo Model

- PK parameters for a modified A1AT have been assessed in vivo
- Statistically significant improvement of PK parameters for modified A1AT compared to Aralast



40 Subject to regulatory approval



WE ARE OPTIMIZING EFFICIENCIES OF PLASMA-DERIVED THERAPY PRODUCTION



Estimated % of
PDT R&D spend for
FY2023



Optimizing value of in-line products



Plasma production efficiencies



New plasma-derived therapies

→ Pharmaceutical science support for manufacturing

41



WE ARE FURTHER IMPROVING MANUFACTURING EFFICIENCIES TO INCREASE YIELD



High yield high throughput initiatives will improve delivery of last liter products to patients globally

A new high yield & high throughput process:

- Process development to shorten IgG upstream and total albumin cycle times
- Capture of purification waste to isolate proteins for possible new development

Potential benefit of higher yield and increased capacity

Significantly reduced COGS with positive ROI



WE ARE IDENTIFYING AND DEVELOPING NEW PLASMA-DERIVED THERAPIES



Estimated % of PDT R&D spend for FY2023



Optimizing value of in-line products



Plasma production efficiencies



New plasma-derived therapies

→ New targeted therapies for diverse therapeutic areas



WE BELIEVE THERE IS A TREMENDOUS AMOUNT OF UNTAPPED POTENTIAL IN PLASMA PROTEINS



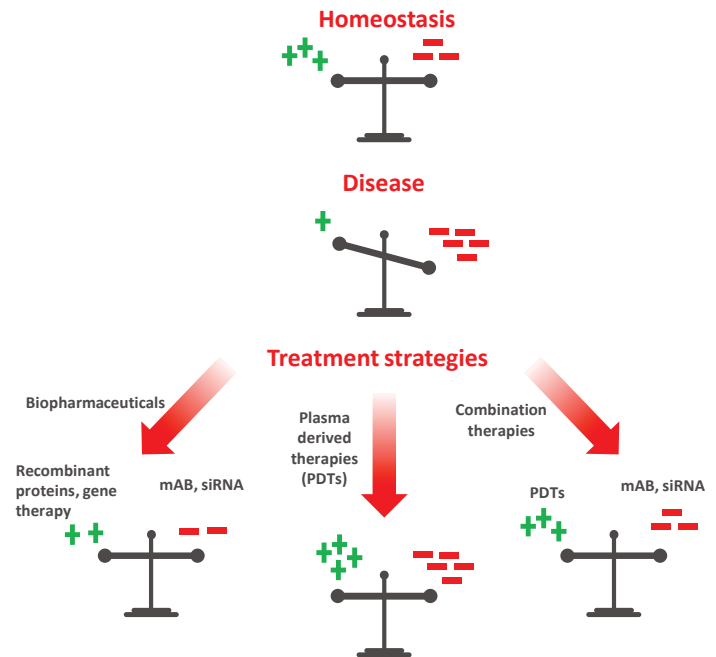
>3000 plasma proteins control balance, some with health promoting + effects and other with disease associated - effects



Generally, PDTs have been developed to **replace functional deficiencies** in health promoting proteins



We believe PDTs, alone or in combination, can be developed to **address acute and chronic diseases**



We are well-positioned to create near-term and sustainable growth



		NEAR TERM CATALYSTS		SUSTAINED GROWTH	
		FY19 – FY22	FY23 – FY24	FY25 AND BEYOND	
TARGET APPROVAL FY					
IMMUNOLOGY	HYQVIA <i>Halozyme</i> Chronic inflammatory demyelinating polyneuropathy (CIDP)		CUVITRU Japan PID (FPI Q4 2019)	GLASSIA <i>Kamada</i> A1ATD-emphysema*	HYPERIMMUNE IGx GENERATION
	GLASSIA <i>Kamada</i> Immunogenicity/bronchioalveolar lavage		HYQVIA <i>Halozyme</i> EU Pediatric PID	CINRYZE Ex-HAE indications TBD	ACUTE PHASE REACTANTS
	HYQVIA - HyHub <i>Flextronics</i> Delivery Device		TAK 880 Low IgA-IgG (IV) Primary Immunodeficiency	CINRYZE Geographic expansion	NEUROIMMUNOLOGY/OTHER AUTOIMMUNE
	HYQVIA Geographic expansion		HYQVIA <i>Halozyme</i> US Pediatric PID	Hyper-Immune IG Infectious disease	PLASMA-DRUG COMBINATIONS
	CUVITRU Geographic expansion		CUVITRU Wearable Device	Alpha-1 Antitrypsin (A1AT) Next generation formulations	INTEGRATED CARE: DEVICES AND DIAGNOSTICS
		TAK 881 Facilitated 20% SC IgG <i>Halozyme</i> Primary Immunodeficiency (PID)			PLASMA PROTEOMICS for BIOMARKERS and NEW DRUG DISCOVERY
HEMATOLOGY	CEPROTIN Geographic expansion		PROTHROMPLEX TOTAL Device and formulation	PROTHROMPLEX TOTAL US - Drug-induced bleeding **	
	FEIBA Volume reduction		Butyryl Cholinesterase Organophosphate poisoning		

*Subject to regulatory approval
**Pending FDA Pre-IND consultation and future acceptance of an IND

TREATMENT PARADIGMS OF RARE AND COMPLEX DISEASES ARE DYNAMIC AND WE ARE INNOVATING CONTINUOUSLY



Uncertainties

PDT Innovation



- Deepening understanding of underlying mechanisms of diseases and co-morbidities

- Directed most appropriate uses of PDTs
- With Takeda Global R&D, investigate plasma-drug combinations



- Evolution of Fc- and Fc-Receptor approaches (including anti-FcRn)
- Gene therapies and RNAi for specific diseases

- Focus on primary and secondary immunodeficiencies
- Identify IG responders in specific auto-immune diseases
- Develop PDTs in conjunction with gene therapies and RNAi (e.g. A1ATD-liver disease)



- Perception of lack of plasma product differentiation

- Integrated care solutions will help to expand therapeutic values and differentiate Takeda products
- New formulations may offer new approaches for patients

46

KEY TAKEAWAYS FOR PLASMA-DERIVED THERAPIES R&D



1

Dedicated PDT R&D organization focused on – and investing in – reimagining plasma, while leveraging Takeda’s broader R&D resources and capabilities

2

Poised to deliver near-term value by optimizing our in-line portfolio and improving efficiencies throughout the value chain

3

Committed to creating long-term value by unlocking the full potential of plasma to develop innovative, integrated solutions that meaningfully benefit patients globally

47



REALIZING THE POTENTIAL OF PLASMA-DERIVED THERAPIES

21st November 2019

Julie Kim

President, Plasma-Derived Therapies Business Unit

