

A solid red vertical bar on the left side of the page.

# **Takeda Quarterly Financial Report**

**For the Quarter Ended December 31, 2021**

---

## Table of Contents

---

<b>Financial Highlights</b>	<b>3</b>
Selected Financial Results	3
Revenue by Region	4
Revenue by Therapeutic Area and Product	5
<b>Recent Developments</b>	<b>7</b>
Business Development	7
Pipeline and R&D Activities	7
<b>Analysis of results of operations, financial position, and cash flow</b>	<b>20</b>
Results of Operations (Reported)	20
Results of Operations (Underlying)	24
Consolidated Financial Position	26
Consolidated Cash Flow	28
Outlook for the Fiscal Year Ending March 31, 2022	29
Impact of the Spread of the Novel Coronavirus Infectious Disease (COVID-19) and Takeda's Initiatives in Response	31
<b>Consolidated Financial Statements</b>	<b>32</b>
(1) Consolidated Statements of Profit or Loss	32
(2) Consolidated Statements of Comprehensive Income	33
(3) Consolidated Statements of Financial Position	34
(4) Consolidated Statements of Changes in Equity	36
(5) Consolidated Statements of Cash Flows	38
(6) Other Information	40
<b>Supplementary Information</b>	<b>41</b>
1. Pipeline	42
• I. Clinical Development Activities	42
• II. Recent Progress in stage	48
• III. Discontinued projects	49
• IV. Main Research & Development collaborations	50
2. Supplementary Financial Information	56
• Revenue by region	56
◦ Year to date	56
◦ Quarterly	57
• Product Sales Analysis	58
◦ Year to date	58
◦ Quarterly	60
■ Q1	60
■ Q2	62
■ Q3	64
• Product Sales Analysis (Reported & Underlying Growth)	66
• Product Forecast	68
• Exchange Rate	70
• CAPEX, depreciation and amortization and impairment losses	71

<u>3. Reconciliation</u> .....	<u>72</u>
• <u>FY2021 Q3 YTD Reconciliation from Reported Revenue to Core/Underlying Revenue</u> .....	<u>72</u>
• <u>FY2021 Q3 YTD Reconciliation from Reported to Core/Underlying Core</u> .....	<u>73</u>
• <u>FY2020 Q3 YTD Reconciliation from Reported to Core/Underlying Core</u> .....	<u>74</u>
• <u>Free Cash Flow</u> .....	<u>75</u>
• <u>FY2021 Q3 YTD LTM Net Profit to Adjusted EBITDA Bridge</u> .....	<u>76</u>
• <u>FY2021 Q3 YTD Net Debt to Adjusted EBITDA</u> .....	<u>77</u>
• <u>FY2020 Net Debt to Adjusted EBITDA</u> .....	<u>78</u>
• <u>Reconciliation from Reported Operating Profit to Core Operating Profit - FY2021 Forecast</u> .....	<u>79</u>
<b><u>Important Notice</u></b> .....	<u>80</u>

## Financial Highlights

### Selected Financial Results

#### Results of Operation

(JPY millions)	December 31,		Change versus the previous year	
	2020	2021	JPY	%
Revenue	2,427,538	2,695,717	268,178	11.0 %
Operating profit	358,729	462,463	103,734	28.9 %
Profit before tax	235,357	356,618	121,261	51.5 %
Net profit for the period	179,027	241,541	62,514	34.9 %
Net profit attributable to owners of the Company	178,907	241,417	62,510	34.9 %
Earnings per share (JPY)				
Basic earnings per share	114.57	154.09	39.52	34.5 %
Diluted earnings per share	113.72	153.03	39.31	34.6 %

#### Non-IFRS Measures

##### Results of Operations

(JPY billions)	December 31,		Change versus the previous year	
	2020	2021	JPY	%
<b>Underlying:</b>				
Revenue Growth	+ 1.1 %	+ 7.1%		
Core operating profit margin	29.9 %	29.4 %		
<b>Core Operating Profit</b>	780.6	757.9	(22.7)	(2.9) %
<b>Core EPS (yen)</b>	333	333	(0)	(0.0) %
<b>Free Cash Flow</b>	717.5	671.3	(46.2)	(6.4) %

##### Leverage

(JPY billions)	As of	
	March 31, 2021	December 31, 2021
<b>Net debt</b>	(3,429.4)	(3,391.9)
<b>Adjusted EBITDA (Last 12 months)</b>	1,083.5	1,144.1
<b>Net debt/Adjusted EBITDA ratio</b>	3.2 x	3.0 x

Takeda uses certain non-IFRS measures to supplement the analysis of results of operations under International Financial Reporting Standards ("IFRS"). Refer to *Supplementary Information "3. Reconciliation"* for reconciliations of non-IFRS Measures.

#### Consolidated Cash Flows

(JPY millions)	December 31,		Change versus the previous year	
	2020	2021	JPY	%
Cash flows from (used in) operating activities	609,971	747,521	137,550	22.6 %
Cash flows from (used in) investing activities	100,199	(172,487)	(272,686)	—
Cash flows from (used in) financing activities	(718,282)	(826,465)	(108,183)	15.1 %

#### Consolidated Financial Position

(JPY millions)	As of		Change versus the previous year	
	March 31, 2021	December 31, 2021	JPY	%
Non-current Assets	10,199,400	10,220,626	21,226	0.2 %
Current Assets	2,712,893	2,477,893	(235,000)	(8.7) %
<b>Total Assets</b>	<b>12,912,293</b>	<b>12,698,519</b>	<b>(213,774)</b>	<b>(1.7) %</b>
Non-current Liabilities	5,961,940	5,557,443	(404,497)	(6.8) %
Current Liabilities	1,773,176	1,809,254	36,078	2.0 %
<b>Total Liabilities</b>	<b>7,735,116</b>	<b>7,366,697</b>	<b>(368,419)</b>	<b>(4.8) %</b>
<b>Equity</b>	<b>5,177,177</b>	<b>5,331,822</b>	<b>154,645</b>	<b>3.0 %</b>
<b>Total liabilities and equity</b>	<b>12,912,293</b>	<b>12,698,519</b>	<b>(213,774)</b>	<b>(1.7) %</b>

**Forecast and Management Guidance**

Forecast\*

(JPY billions)	Previous Forecast (October 28, 2021)	Revised Forecast (February 3, 2022)	vs. Previous Forecast	
<b>Reported:</b>				
Revenue	3,370.0	3,510.0	140.0	4.2 %
Operating profit	488.0	515.0	27.0	5.5 %
Profit before tax	352.0	385.0	33.0	9.4 %
Net profit for the year (attributable to owners of the Company)	184.3	242.5	58.2	31.6 %
EPS (JPY)	117.35	154.99	37.64	32.1 %
<b>Non-IFRS Measures</b>				
Core Operating Profit	930.0	970.0	40.0	4.3 %
Core EPS (JPY)	394	416	22	5.6 %
Free cash flow (including announced divestitures)	600.0 - 700.0	700.0 - 800.0		
<b>Dividends per share (Yen)</b>	180	180	—	— %

\*Refer to *Analysis of Results of Operations, Financial Position, and Cash Flow* "[Outlook for the Fiscal Year Ending March 31, 2022](#)" for details.

Management Guidance\*

	Guidance as of October 28, 2021	Guidance as of February 3, 2022
Underlying Revenue Growth	Mid-single-digit growth	Mid-single-digit growth
Underlying Core Operating Profit Growth	Mid-single-digit growth	Mid-single-digit growth
Underlying Core Operating Profit Margin	~30% margin	~30% margin
Underlying Core EPS Growth	Mid-single-digit growth	Mid-single-digit growth

\*Underlying growth adjusts for divestitures (assets divested in FY2020 and disclosed divestitures expected to close in FY2021) and applies a constant exchange rate. Please refer to *Analysis of Results of Operations, Financial Position, and Cash Flow* "[Results of Operations \(Underlying\)](#)" for definition of underlying growth.

**Revenue by Region**

		JPY (millions)							
		Nine-month Period Ended December 31,							
		Japan	United States	Europe and Canada	Asia (excluding Japan)	Latin America	Russia/CIS	Other	Total
	2020	435,112	1,188,965	499,962	119,178	95,414	38,724	50,183	2,427,538
	2021	530,245	1,297,020	540,978	139,770	93,545	43,582	50,577	2,695,717
Change versus the previous year	JPY	95,133	108,055	41,016	20,592	(1,868)	4,858	393	268,178
	%	21.9 %	9.1 %	8.2 %	17.3 %	(2.0)%	12.5 %	0.8 %	11.0 %

“Other” includes the Middle East, Oceania and Africa. This disaggregation provides revenue attributable to countries or regions based on the customer location.

## Revenue by Therapeutic Area and Product

	JPY (millions)			
	Nine-month Period Ended December 31,		Change versus the previous year	
	2020	2021	JPY	%
<b>Gastroenterology:</b>				
ENTYVIO	319,307	395,373	76,066	23.8 %
TAKECAB-F <sup>(1)</sup>	64,134	78,373	14,239	22.2 %
GATTEX/REVESTIVE	50,149	56,635	6,486	12.9 %
DEXILANT	43,458	40,136	(3,322)	(7.6)%
PANTOLOC/CONTROLOC <sup>(2)</sup>	32,383	30,068	(2,314)	(7.1)%
LIALDA/MEZAVANT	18,725	19,042	317	1.7 %
PENTASA	17,833	15,771	(2,062)	(11.6)%
AMITIZA	18,826	5,865	(12,961)	(68.8)%
RESOLOR/MOTEGRITY	8,517	10,100	1,583	18.6 %
ALOFISEL	565	1,353	788	139.5 %
Others	14,914	12,967	(1,948)	(13.1)%
<b>Total Gastroenterology</b>	<b>588,811</b>	<b>665,683</b>	<b>76,871</b>	<b>13.1 %</b>
<b>Rare Diseases:</b>				
Rare Metabolic:				
ELAPRASE	51,531	57,714	6,183	12.0 %
REPLAGAL	38,874	39,568	693	1.8 %
VPRIV	28,868	32,171	3,303	11.4 %
NATPARA/NATPAR	2,503	3,926	1,423	56.9 %
Total Rare Metabolic	121,776	133,378	11,602	9.5 %
Rare Hematology:				
ADVATE	97,112	89,315	(7,796)	(8.0)%
ADYNOVATE/ADYNOVI	43,765	45,873	2,108	4.8 %
FEIBA	34,235	28,978	(5,257)	(15.4)%
RECOMBINATE	10,457	9,586	(871)	(8.3)%
HEMOFIL/IMMUNATE/IMMUNINE	13,233	13,543	309	2.3 %
Other PDT Products	2,580	3,032	451	17.5 %
Others	17,192	21,266	4,074	23.7 %
Total Rare Hematology	218,574	211,592	(6,982)	(3.2)%
Hereditary Angioedema:				
TAKHZYRO	65,891	78,425	12,534	19.0 %
FIRAZYR	20,100	21,471	1,371	6.8 %
CINRYZE	17,264	14,703	(2,560)	(14.8)%
KALBITOR	3,103	3,139	36	1.2 %
Total Hereditary Angioedema	106,357	117,738	11,381	10.7 %
Others:	—	190	190	-
<b>Total Rare Diseases</b>	<b>446,707</b>	<b>462,897</b>	<b>16,190</b>	<b>3.6 %</b>
<b>PDT Immunology:</b>				
immunoglobulin	248,031	278,309	30,278	12.2 %
albumin	43,599	61,490	17,891	41.0 %
Other	21,410	23,448	2,039	9.5 %
<b>Total PDT Immunology</b>	<b>313,040</b>	<b>363,247</b>	<b>50,208</b>	<b>16.0 %</b>

JPY (millions)

	Nine-month Period Ended December 31,		Change versus the previous year	
	2020	2021	JPY	%
<b>Oncology:</b>				
VELCADE	75,892	84,459	8,567	11.3 %
LEUPLIN/ENANTONE	75,255	82,215	6,960	9.2 %
NINLARO	67,863	70,747	2,884	4.2 %
ADCETRIS	44,385	51,786	7,402	16.7 %
ICLUSIG	26,259	26,687	428	1.6 %
VECTIBIX	18,376	19,355	979	5.3 %
ALUNBRIG	6,483	10,127	3,644	56.2 %
Other	3,949	13,720	9,771	247.4 %
<b>Total Oncology</b>	<b>318,462</b>	<b>359,096</b>	<b>40,635</b>	<b>12.8 %</b>
<b>Neuroscience:</b>				
VYVANSE/ELVANSE	202,430	244,994	42,564	21.0 %
TRINTELLIX	52,680	63,030	10,350	19.6 %
INTUNIV	14,851	12,525	(2,326)	(15.7) %
ADDERALL XR	13,353	15,957	2,603	19.5 %
ROZEREM	9,500	9,424	(76)	(0.8) %
Other	22,285	16,701	(5,583)	(25.1) %
<b>Total Neuroscience</b>	<b>315,098</b>	<b>362,630</b>	<b>47,532</b>	<b>15.1 %</b>
<b>Other:</b>				
AZILVA-F <sup>(1)</sup>	62,793	60,057	(2,736)	(4.4) %
LOTRIGA	24,466	24,753	287	1.2 %
AIPHAGAN	12,228	12,000	(229)	-1.9 %
FOSRENOL	10,208	10,187	(21)	-0.2 %
ACTOVEGIN	8,276	11,046	2,770	33.5 %
Others <sup>(3)</sup>	327,449	364,121	36,672	11.2 %
<b>Total Other</b>	<b>445,420</b>	<b>482,163</b>	<b>36,743</b>	<b>8.2 %</b>
<b>Total Revenue by Product</b>	<b>2,427,538</b>	<b>2,695,717</b>	<b>268,178</b>	<b>11.0 %</b>

<sup>(1)</sup> The figures include the amounts of fixed dose combinations and blister packs.

<sup>(2)</sup> Generic name: pantoprazole

<sup>(3)</sup> The figure for the nine-month period ended December 31, 2020 includes the revenue of Takeda Consumer Healthcare Company Limited, which was divested on March 31, 2021. The figure for the nine-month period ended December 31, 2021 includes the 133,043 million JPY selling price on sales of four diabetes products (NESINA, LIOVEL, INISYNC and ZAFATEK) in Japan to Teijin Pharma Limited recorded as revenue.

## Recent Developments

### **Business Development**

During the three-month period ended December 31, 2021 and through the issuance of its earnings release dated February 3, 2022, Takeda Pharmaceutical Company Limited ("Takeda", or the "Company") divested certain businesses and assets in non-core areas as part of its efforts to deleverage toward its target of 2x (i.e. "low-twos") net debt/adjusted EBITDA within March 2022 - March 2024. Major divestment activities during the period are as follows:

- In April 2021, we completed the asset transfer associated with a portfolio of select non-core products in Japan to Teijin Pharma Limited for a total value of 133.0 billion JPY.

### **Pipeline and R&D Activities**

Research and development expenses for the nine-month period ended December 31, 2021 were 382.5 billion JPY.

Takeda's R&D engine is focused on translating science into highly innovative, life-changing medicines that make a critical difference to patients. Takeda supports dedicated R&D efforts across three areas: Innovative Biopharma, Plasma-Derived Therapies (PDT) and Vaccines. The R&D engine for Innovative Biopharma is the largest component of our R&D investment and has produced exciting new molecular entities (NMEs) that represent potential best-in-class and/or first-in-class medicines in areas of high unmet medical need across our core Therapeutic Areas (oncology, rare genetics and hematology, neuroscience, and gastroenterology (GI)). Over the past several years, and more recently bolstered by our acquisition of Shire, we have also harnessed the potential of cell and gene therapies by investing in new capabilities and next-generation platforms internally and through a network of partnerships.

Takeda's pipeline is positioned to support both near-term and sustained growth of the company. Once first approval is achieved, there is ongoing R&D support for geographical expansion and additional indications, as well as post-marketing commitment and potential additional formulation work. Takeda's R&D team works closely with the commercial functions to maximize the value of marketed products and reflect commercial insights in its R&D strategies and portfolio.

Major progress on R&D events since April 2021 are listed as follows:

### **R&D pipeline**

#### **Oncology**

In Oncology, Takeda endeavors to deliver novel medicines to patients with cancer worldwide through a commitment to breakthrough innovation and a passion for improving the lives of patients. Takeda focuses on three key areas in oncology: (1) building on its foundational expertise in hematologic malignancies through continued investment in lifecycle management programs for marketed products NINLARO, ADCETRIS, and ICLUSIG, as well as in pipeline assets in Multiple Myeloma, Acute Myeloid Leukemia, Myelodysplastic Syndromes, and other blood cancers; (2) further developing its portfolio in lung cancer with the marketed products ALUNBRIG, EXKIVITY, and development programs in targeted lung cancer populations; and (3) pursuing novel immuno-oncology targets and next-generation platforms, as well as exploring innovative cell therapies harnessing the power of the innate immune system.



*NINLARO / Generic name: ixazomib*

- In May 2021, Takeda announced that it received approval from the Japanese Ministry of Health, Labour and Welfare (MHLW) for a partial amendment to the manufacturing and marketing approval of NINLARO to expand the eligible patient population for this medicine to those requiring a maintenance therapy after first-line treatment for multiple myeloma without prior stem cell transplant. The approval is based primarily on the results of the TOURMALINE-MM4 study, a randomized and placebo-controlled double-blind multicenter international Phase III clinical trial. The study achieved its primary endpoint, demonstrating a statistically significant improvement in progression-free survival (PFS) in adult patients with multiple myeloma receiving NINLARO maintenance who had not undergone stem cell transplantation. The safety profile of NINLARO as a maintenance therapy is similar to its established safety profile in the monotherapy setting, and, notably, no new concerns were identified in the TOURMALINE-MM4 study.

*ICLUSIG / Generic name: ponatinib*

- In June 2021, Takeda presented primary analysis data from the Phase II OPTIC (Optimizing Ponatinib Treatment in CML) trial during an oral session at the virtual 57<sup>th</sup> American Society of Clinical Oncology (ASCO) Annual Meeting, and as an oral session at the virtual 26<sup>th</sup> European Hematology Association (EHA) Annual Meeting. The OPTIC trial, which evaluated treatment in patients with resistant disease, with and without mutations, met its primary endpoint. The study demonstrated that the optimal benefit-risk profile for ICLUSIG in patients with CP-CML is achieved with a daily starting dose of 45-mg and, upon achieving  $\leq 1\%$  BCR-ABL1<sup>IS</sup>, dose reduction to 15-mg. The results also suggest a clinically manageable safety and arterial occlusive event (AOE) profile for ICLUSIG.

*ALUNBRIG / Generic name: brigatinib*

- In June 2021, Takeda announced that ALUNBRIG can be used for first-line treatment of patients with non-small cell lung cancer (NSCLC) who are ALK fusion gene positive (ALK-positive) as determined by the companion diagnostic ALK fusion protein kit, Ventana OptiView ALK (D5F3) (Ventana) in Japan. Ventana, developed by Roche Diagnostics, which uses as its assay principle the immunohistochemical staining method (IHC method), received an additional indication through a partial change of the drug's manufacturing and marketing approval to include its use to ALUNBRIG. The additional approval of ALUNBRIG for the indication of Ventana, in addition to the Fluorescence *In Situ* Hybridization (FISH) diagnostic, will provide a wider range of ALK-positive NSCLC patients with the opportunity to be treated with ALUNBRIG.

*ADCETRIS / Generic name: brentuximab vedotin*

- In September 2021, Takeda announced that it submitted a Supplemental New Drug Application (sNDA) of ADCETRIS in the first-line treatment of CD30-positive Hodgkin lymphoma in pediatric patients in Japan. This application is based on the results of a global Phase 1/2 trial (C25004 Trial) evaluating the efficacy and safety of ADCETRIS in combination with AVD (doxorubicin, vinblastine and dacarbazine) as a first-line therapy in pediatric patients with previously untreated advanced-stage Hodgkin lymphoma.

*CABOMETYX / Generic name: cabozantinib*

- In August 2021, Takeda and Ono Pharmaceutical (Ono) announced that the companies received an approval from the Japanese Ministry of Health, Labour and Welfare (MHLW) for CABOMETYX and Ono's OPDIVO (nivolumab), a human anti-human PD-1 monoclonal antibody, in combination therapy for the treatment of unresectable or metastatic renal cell carcinoma (RCC), for a partial change in approved items of the manufacturing and marketing approval. This approval is based on results from the global, multi-center, randomized, open-label Phase 3 CheckMate-9ER study, evaluating OPDIVO and CABOMETYX combination therapy versus sunitinib alone in patients with previously untreated advanced or metastatic RCC. In this study, OPDIVO and CABOMETYX combination therapy demonstrated a significant and clinically meaningful improvement in the primary endpoint of progression-free survival (PFS) as assessed by the blind independent central review (BICR), compared to sunitinib alone at the final analysis, as well as the secondary endpoints of overall survival (OS) and objective response rate (ORR) as assessed by the BICR. The safety profiles of OPDIVO and CABOMETYX combination therapy observed in the study were consistent with the previously reported safety profile of each product.

*ZEJULA / Generic name: niraparib*

- In September 2021, Takeda announced that it has received approval from the Japanese Ministry of Health, Labour and Welfare (MHLW) to manufacture and market ZEJULA tablets 100mg (ZEJULA tablets) as an additional formulation for ZEJULA capsules 100mg (ZEJULA capsules), an oral poly (ADP-ribose) polymerase (PARP) inhibitor. The approval was granted based on the results of a human bioequivalence trial (3000-01-004 trial) and an dissolution study that confirmed the equivalence of ZEJULA capsules and ZEJULA tablets. ZEJULA capsules require refrigerated storage, however the newly approved ZEJULA tablets can be stored at room temperature.

*EXKIVITY / Generic name: mobocertinib*

- In May 2021, Takeda announced updated data from the Phase 1/2 trial of mobocertinib in patients with epidermal growth factor receptor (EGFR) Exon20 insertion mutation-positive (insertion+) metastatic non-small cell lung cancer (mNSCLC) who received prior platinum-based chemotherapy. The results showed mobocertinib continued to demonstrate clinically meaningful benefit after over a year of follow up and were presented at the virtual 57th American Society of Clinical Oncology (ASCO) Annual Meeting. Results showed a median overall survival (OS) of 24 months with a median follow up of 14 months, and responses were observed across diverse EGFR Exon20 insertion variants. Other key data points such as confirmed objective response rate (ORR), a median duration of response (DoR) and a disease control rate (DCR), remained consistent with previously reported data. The safety profile observed was manageable and consistent with previous findings.
- In July 2021, Takeda announced that Center for Drug Evaluation (CDE) of the National Medical Products Administration of China (NMPA) has accepted the New Drug Application (NDA) for mobocertinib and granted priority review for this Class-1 innovative drug, for the treatment of adult patients with non-small cell lung cancer (NSCLC) with epidermal growth factor receptor (EGFR) Exon20 insertion mutations.

- In September 2021, Takeda announced that the U.S. Food and Drug Administration (FDA) has approved EXKIVITY for the treatment of adult patients with locally advanced or metastatic non-small cell lung cancer (NSCLC) with epidermal growth factor receptor (EGFR) exon 20 insertion mutations as detected by an FDA-approved test, whose disease has progressed on or after platinum-based chemotherapy. The FDA approval is based on results from the platinum-pretreated population in the Phase 1/2 trial of EXKIVITY, which consisted of 114 patients with EGFR Exon20 insertion+ NSCLC who received prior platinum-based therapy and were treated at the 160 mg dose once- daily. EXKIVITY, which was granted priority review and received Breakthrough Therapy Designation, Fast Track Designation and Orphan Drug Designation from the FDA, is the first and only approved oral therapy specifically designed to target EGFR Exon20 insertion mutations. This indication is approved under Accelerated Approval based on overall response rate (ORR) and duration of response (DoR). Continued approval for this indication may be contingent upon verification and description of clinical benefit in a confirmatory trial. The FDA simultaneously approved Thermo Fisher Scientific's OncoPrint Dx Target Test as a next-generation sequencing (NGS) companion diagnostic for EXKIVITY to identify NSCLC patients with EGFR Exon20 insertions.

*Development code: TAK-924 / Generic name: pevonedistat*

- In September 2021, Takeda announced the Phase 3 PANTHER (Pevonedistat-3001) study did not achieve pre-defined statistical significance for the primary endpoint of event-free survival (EFS). The trial evaluated whether the combination of pevonedistat plus azacitidine as first-line treatment for patients with higher-risk myelodysplastic syndromes (MDS), chronic myelomonocytic leukemia (CMML) and low-blast acute myeloid leukemia (AML) improved EFS versus azacitidine alone. An event in the trial was defined as death or transformation to AML in participants with higher-risk MDS or CMML, whichever occurred first, and death in participants with AML. Takeda discontinued all research and development.

**Rare Genetics & Hematology**

In Rare Genetics & Hematology, Takeda focuses on several areas of high unmet medical need. In hereditary angioedema, Takeda aspires to transform the treatment paradigm, including through TAKHZYRO, with continued investment in lifecycle management programs including evaluating Takhzyro in Bradykinin-mediated angioedema with normal C1-inhibitor. In rare hematology, Takeda focuses on addressing today's needs in the treatment of bleeding disorders, including through ADVATE and ADYNOVATE/ADYNOVI, as well as on the development of pipeline assets including TAK-755 for the treatment of immune thrombotic thrombocytopenic purpura (iTTP) and congenital thrombotic thrombocytopenic purpura (cTTP). In rare metabolic diseases, Takeda is developing treatments for lysosomal storage disorders (LSDs), with a portfolio that includes commercial products such as ELAPRASE and REPLAGAL, and late-stage investigational therapies and pipeline candidates. We are also building differentiated gene therapy capabilities for the development and delivery of functional cures to patients with rare diseases.

*TAKHZYRO / Generic name: lanadelumab*

- In July 2021, Takeda announced the results from two final analyses from the Phase 3 HELP (Hereditary Angioedema Long-term Prophylaxis) Study™ Open-label Extension (OLE), which evaluated the long-term safety (primary endpoint) and efficacy of TAKHZYRO (lanadelumab) 300 mg every two weeks for up to 2.5 years. In the first analysis, the mean (min, max) reduction in the attack rate compared to baseline observed in the study population (N=212) was of 87.4 percent (-100; 852.8), and the median reduction was 97.7 percent and patients

received treatment for a mean (standard deviation) duration of 29.6 (8.2) months. At steady state – day 70 to the end of the treatment period – attack rates were further reduced to a mean of 92.4 percent and a median reduction of 98.2 percent. An additional analysis further suggests TAKHZYRO was a well-tolerated treatment that prevented HAE attacks over an extended planned 132 week treatment period across specific HAE patient demographic and disease characteristic subgroups. These data were presented at the 2021 European Academy of Allergy and Clinical Immunology (EAACI) Hybrid Congress.

*REPLAGAL / Generic name: agalsidase alfa*

- In November 2021, Takeda and Sumitomo Dainippon Pharma Co., Ltd. (Sumitomo Dainippon Pharma) announced that Takeda will assume the manufacturing and marketing authorization (and the marketing rights) of REPLAGAL 3.5mg for Fabry disease, an  $\alpha$ -galactosidase enzyme intravenous (IV) infusion, from Sumitomo Dainippon Pharma as of February 15, 2022 in Japan.

*FIRAZYR / Generic name: icatibant*

- In December 2021, Takeda announced that it has submitted an application for a revision to the marketing approval for the selective bradykinin B2 receptor blocker FIRAZYR for the treatment of pediatric patients with hereditary angioedema (HAE) in Japan. This application is based primarily on a Japanese Phase III open-label study and an overseas Phase III open-label study evaluating the safety, efficacy and pharmacokinetics of subcutaneous administration of FIRAZYR in children mainly aged between two and 18 years. The Japanese pediatric treatment response in the Japanese Phase III open-label study was similar to the pediatric treatment response in Japanese and overseas adults and in the overseas Phase III open-label study.

*VONVENDI / Generic name: von Willebrand factor (Recombinant)*

- In January 2022, Takeda announced that the U.S. Food & Drug Administration (FDA) approved VONVENDI for routine prophylaxis to reduce the frequency of bleeding episodes in patients with severe Type 3 von Willebrand disease (VWD) receiving on-demand therapy. The approval is based on data from a prospective, open-label, international multicenter study to evaluate efficacy and safety of prophylactic treatment of VONVENDI in reducing the frequency of bleeding episodes in 10 adult patients diagnosed with severe Type 3 VWD who were previously treated on-demand. VONVENDI is now indicated for routine prophylaxis in adults with severe Type 3 VWD receiving on-demand therapy, as well as on-demand and perioperative bleed management in adults with VWD.

*LIVTENCITY / Generic name: maribavir*

- In May 2021, Takeda announced that the U.S. Food and Drug Administration (FDA) accepted a New Drug Application (NDA), granting priority review, for maribavir for the treatment of CMV infections that are refractory with or without resistance (R/R), in solid organ transplant (SOT) or hematopoietic cell transplant (HCT) recipients. The application is based on the pivotal Phase 3 TAK-620-303 (SOLSTICE) trial. Maribavir has been granted Orphan Drug Designation by the FDA for treatment of clinically significant CMV viremia and disease in at-risk patients. The FDA has also granted maribavir Breakthrough Therapy Designation as a treatment for CMV infection and disease in transplant patients resistant or refractory to prior therapy.

- In June 2021, Takeda announced the results from a new subgroup analysis of SOT recipients in the Phase 3 TAK-620-303 (SOLSTICE) trial, for the investigational drug maribavir, at the American Transplant Congress (ATC) 2021 Virtual Connect. More than twice (55.6%, 79/142) as many SOT recipients with R/R CMV infection at baseline treated with maribavir achieved confirmed CMV viremia clearance at Study Week 8 (end of treatment phase) compared to those treated with conventional antiviral therapies (26.1%, 18/69) (investigator assigned treatment; IAT consists of one or a combination of ganciclovir, valganciclovir, foscarnet or cidofovir) (adjusted difference [95% CI]: 30.5% [17.3, 43.6]). The results presented showed consistent efficacy in SOT recipients receiving maribavir in heart, lung and kidney transplants.
- In October 2021, Takeda announced that the U.S. Food and Drug Administration (FDA) Antimicrobial Drugs Advisory Committee (AMDAC) voted unanimously to recommend use of maribavir for the treatment of refractory cytomegalovirus (CMV) infection and disease with genotypic resistance to ganciclovir, valganciclovir, foscarnet or cidofovir in transplant recipients. The committee also voted unanimously to recommend use of maribavir for the treatment of refractory CMV infection and disease without genotypic resistance to ganciclovir, valganciclovir, foscarnet or cidofovir in transplant recipients. Both recommendations were based on the results of the Phase 2 and Phase 3 TAK-620-303 (SOLSTICE) trials. The New Drug Application (NDA) for maribavir is currently under Priority Review by the FDA. The FDA will consider the vote as part of its review of the NDA and is not bound by the AMDAC's recommendation.
- In November 2021, Takeda announced that the U.S. Food and Drug Administration (FDA) approved LIVTENCITY for the treatment of adults and pediatric patients (12 years of age or older and weighing at least 35 kg) with post-transplant cytomegalovirus (CMV) infection/disease that is refractory to treatment (with or without genotypic resistance) with ganciclovir, valganciclovir, cidofovir, or foscarnet. Prior to FDA approval, LIVTENCITY was granted Orphan Drug Designation by the FDA for treatment of clinically significant CMV viremia and disease in at-risk patients, as well as Breakthrough Therapy Designation as a treatment for CMV infection and disease in transplant patients resistant or refractory to prior therapy. Takeda is also investigating LIVTENCITY as a first-line treatment of CMV in hematopoietic stem cell transplant recipients in an ongoing Phase 3 clinical trial.
- In December 2021, Takeda announced that the data from the pivotal Phase 3 SOLSTICE clinical trial of LIVTENCITY in post-transplant refractory CMV infections with or without resistance (R/R) were published in the journal of Clinical Infectious Diseases. The SOLSTICE study primary endpoint was met, with 55.7% (131/235) of adult patients on LIVTENCITY achieving confirmed CMV DNA level below the lower limit of quantification (<LLOQ, i.e. <137 IU/mL) at the end of Study Week 8 (end of treatment phase) in comparison with 23.9% (28/117) of patients on conventional antiviral therapies (one or a combination of ganciclovir, valganciclovir, foscarnet or cidofovir); adjusted difference [95% CI]: 32.8% [22.80 to 42.74]; P<0.001. The key secondary endpoint of the composite achievement of CMV DNA level <LLOQ and symptom control at Week 8 maintained through Week 16 was met, with a higher proportion of patients in the LIVTENCITY arm (18.7%, 44/235) meeting the endpoint compared to those on conventional antiviral therapies (10.3%, 12/117); adjusted difference [95% CI]: 9.5% [2.02 to 16.88]; P=0.013.

## Neuroscience

In Neuroscience, Takeda is focusing its R&D investments on potentially transformative treatments for neurological and neuromuscular diseases of high unmet need, and building its pipeline through a combination of in-house expertise and partnerships. By harnessing advances in disease biology understanding, translational tools, and innovative modalities, Takeda is primarily focusing on rare neurology, in particular, on potential investigative therapies for sleep-wake disorders such as narcolepsy and idiopathic hypersomnia with a franchise of orexin-2 receptor agonists (TAK-994, TAK-861, TAK-925, etc.), and rare epilepsies with soticlestat (TAK-935). Other rare neurology diseases of focus include Amyotrophic Lateral Sclerosis, Huntington's disease and other ataxias. Takeda also makes targeted investments to potentially address well-defined segments of neurodegenerative diseases (e.g., Parkinson's Disease).

### *Development code: TAK-994*

- In July 2021, Takeda announced that the U.S. Food and Drug Administration (FDA) granted Breakthrough Therapy Designation (BTD) to TAK-994, its Phase 2 investigational oral orexin agonist, which is designed to selectively target orexin 2 receptors. TAK-994 is currently being studied in an ongoing Phase 2 (TAK-994-1501) study for the treatment of excessive daytime sleepiness (EDS) in patients with narcolepsy type 1 (NT1), a chronic neurological disorder that alters the sleep-wake cycle. The TAK-994 BTD was based, in part, on early phase and preliminary clinical data that indicates Takeda's investigational oral orexin agonist may demonstrate substantially improved objective and subjective measurements of daytime wakefulness in NT1 patients.
- In October 2021, Takeda announced that a safety signal has emerged in Phase 2 studies of TAK-994 (TAK-994-1501 study and TAK-994-1504 study). As an immediate precautionary measure, Takeda has suspended dosing of patients and has decided to stop both Phase 2 studies early. This allows for the timely interpretation of the benefit/risk profile of TAK-994 and to determine next steps for the program.

## Gastroenterology (GI)

In Gastroenterology, Takeda focuses on delivering innovative, life-changing therapeutics for patients with gastrointestinal and liver diseases. Takeda is maximizing the potential of our inflammatory bowel disease (IBD) franchise around ENTYVIO, including development of a subcutaneous formulation as well as a needle free device. Takeda is also expanding its position with GATTEX / REVESTIVE, and ALOFISEL, which is in ongoing P-3 trials to support further potential geographic expansion, including in the U.S. Furthermore, Takeda is progressing a pipeline built through partnerships exploring opportunities in IBD, celiac disease, select liver diseases, and motility disorders.

### *ENTYVIO / Generic name: vedolizumab*

- In October 2021, Takeda announced the update on the U.S. development program for the investigational subcutaneous (SC) formulation of ENTYVIO as a maintenance therapy in adults with moderate to severe ulcerative colitis (UC). Through our ongoing interactions with the U.S. Food and Drug Administration (FDA), Takeda has received feedback which has provided clarity on the regulatory package and critical elements for the resubmission of the Biologics License Application (BLA) for Entyvio SC, and we are moving forward accordingly. We are reviewing our development program timelines and currently anticipate potential approval in FY 2023.
- In December 2021, Takeda announced that the Committee for Medicinal Products for Human Use (CHMP) of the European Medicines Agency (EMA) has recommended the approval of intravenous (IV) ENTYVIO for the treatment of adult patients with moderately to severely active chronic pouchitis, who have undergone proctocolectomy and ileal



pouch-anal anastomosis for ulcerative colitis, and have had an inadequate response with or lost response to antibiotic therapy. The positive opinion from the CHMP was based on the EARNEST trial, recently presented at the United European Gastroenterology's annual meeting, UEG Week Virtual 2021, which assessed the safety and efficacy of ENTYVIO IV in the treatment of active chronic pouchitis. Moreover, information from a number of retrospective studies of historical data indicating that ENTYVIO can have a positive impact on patients with inflammation of the pouch was also included in the application. In January 2022, European Commission (EC) approved ENTYVIO as the first treatment indicated for active chronic pouchitis across the European Union.

*GATTEX / REVESTIVE / Generic name: teduglutide*

- In June 2021, Takeda announced that it obtained approval from the Japanese Ministry of Health, Labour and Welfare (MHLW) to manufacture and market REVESTIVE 3.8 mg for subcutaneous injection as a treatment for short bowel syndrome. The approval is mainly based on the results of several trials conducted overseas, as well as Phase 3 clinical trials (SHP633-302, SHP633-305, SHP633-306, and SHP633-307) conducted in pediatric and adult patients in Japan.
- In November 2021, Takeda announced that it submitted the New Drug Application to the Japanese Ministry of Health, Labour and Welfare (MHLW) for the low dose formulation (0.95 mg) as an additional dosage for REVESTIVE as a treatment for short bowel syndrome (SBS). This new formulation would allow REVESTIVE to be administered to SBS patients weighing less than 10 kg, or less than 20 kg with moderate or severe renal impairment (creatinine clearance of less than 50 mL/min), who cannot be dosed with the 3.8 mg formulation.

*ALOFISEL / Generic name: darvadstrocel*

- In September 2021, Takeda announced that it has received approval from the Japanese Ministry of Health, Labour and Welfare (MHLW) to manufacture and market ALOFISEL for the treatment of complex perianal fistulas in patients with non-active or mildly active luminal Crohn's disease (CD). This product is indicated for the treatment of patients who have shown an inadequate response to at least one existing medicinal treatment. The approval is based on data from two trials, the Japanese Study Darvadstrocel-3002 and the ADMIRE-CD trial, conducted in Europe and Israel. ALOFISEL is the first expanded human allogeneic adipose-derived mesenchymal stem cell therapy to be approved in Japan, which exhibits immunomodulatory and local anti-inflammatory effects at the site of inflammation.

*Development code: TAK-721 (Planned trade name: Eohilia) / Generic name: budesonide oral suspension*

- In December 2021, Takeda announced that it has received a Complete Response Letter (CRL) from the U.S. Food and Drug Administration (FDA) in response to its New Drug Application (NDA) for TAK-721 for the treatment of eosinophilic esophagitis, a chronic inflammatory disease of the esophagus. The CRL indicates the FDA has completed its review of the TAK-721 NDA and determined that it cannot be approved in its present form. In addition, the FDA recommended an additional clinical study in order to help resolve FDA feedback. Takeda announced the discontinuation of this program in February 2022.

## **Plasma-Derived Therapies (PDT)**

Takeda created a dedicated PDT business unit with a focus to manage the business end-to-end, from plasma collection to manufacturing, R&D, and commercialization. In PDT, we aspire to develop life-saving plasma derived treatments which are essential for patients with a variety of rare and complex chronic diseases. The dedicated R&D organization in PDT is charged with maximizing the value of existing therapies, identifying new targeted therapies, and optimizing efficiencies of current product manufacturing. Near-term, our priority is focused on delivering value from our broad immunoglobulin portfolio (HYQVIA, CUVITRU, GAMMAGARD and GAMMAGARD S/D) through pursuit of new indications, geographic expansions, and enhanced patient experience through integrated healthcare technologies. In our hematology and specialty care portfolio, our priority is pursuing new indication and formulation development opportunities for PROTHROMPLEX (4F-PCC), FEIBA, CEPROTIN and ARALAST. Additionally, we are developing next generation immunoglobulin products with 20% fSCIg (TAK-881), IgG Low IgA (TAK-880) and pursuing other early stage opportunities that would add to our diversified commercial portfolio of more than 20 therapeutic products distributed worldwide.

*Development code: CoVIg-19 (previously TAK-888) / Generic name: anti-SARS-CoV-2 polyclonal hyperimmune immunoglobulin*

- In April 2021, The CoVIg-19 Plasma Alliance announced that the Phase 3 Inpatient Treatment with Anti-Coronavirus Immunoglobulin (ITAC) clinical trial sponsored and funded by the National Institute of Allergy and Infectious Diseases (NIAID), part of the National Institutes of Health (NIH), did not meet its endpoints. No serious safety signals were raised in the trial. The study aimed to determine whether an investigational anti-coronavirus hyperimmune intravenous immunoglobulin (H-Ig) medicine could reduce the risk of disease progression when added to standard of care treatment including remdesivir in hospitalized adult patients at risk for serious complications. Analyses remain ongoing and NIAID and the INSIGHT Network intend to publish the full results of the trial soon. Following the outcome of the ITAC trial, the CoVIg-19 Plasma Alliance's work has now concluded.

## **Vaccines**

In Vaccines, Takeda is applying innovation to tackle some of the world's most challenging infectious diseases such as dengue, COVID-19, zika, and norovirus. To support the expansion of our pipeline and the development of our programs, we have entered into partnerships with government organizations in Japan and the U.S., and leading global institutions. Such partnerships have been essential in building the critical capabilities that will be necessary to deliver on our programs and realize their full potential.

*SPIKEVAX (formerly COVID-19 Vaccine Moderna) Intramuscular Injection / Development code: mRNA-1273 (Japanese development code: TAK-919)*

- In May 2021, Takeda announced positive interim results from the ongoing Phase 1/2 immunogenicity and safety clinical trial of TAK-919 in Japan have been submitted to the Japan Pharmaceuticals and Medical Devices Agency (PMDA). Takeda currently has a three-way agreement with Moderna, Inc. (Moderna) and the Government of Japan's Ministry of Health Labour and Welfare (MHLW) to import and distribute 50 million doses of TAK-919 in Japan. This interim analysis showed binding antibody and neutralizing antibody titres were elevated at 28 days after the second dose in 100% of people vaccinated with two 0.5ml doses of TAK-919 given 28 days apart. The vaccine candidate was generally well-tolerated with no significant safety concerns reported. The study results were submitted to the Japan Pharmaceuticals and Medical Devices Agency (PMDA) to be evaluated as part of the New



Drug Application submitted in March 2021, which also includes safety and efficacy results from Moderna's pivotal Phase 3 COVE trial conducted in the U.S.

- In May 2021, Takeda announced that the Japanese Ministry of Health, Labour and Welfare (MHLW) granted special approval under article 14-3 of the Pharmaceuticals and Medical Devices Act for emergency use of COVID-19 Vaccine Moderna Intramuscular Injection in Japan. The approval is based on positive clinical data from Takeda's Phase 1/2 immunogenicity and safety clinical trial of COVID-19 Vaccine Moderna Intramuscular Injection in Japan, which showed an immune response consistent with results from Moderna's pivotal Phase 3 COVE trial conducted in the United States. Takeda has started distribution in Japan.
- In July 2021, Takeda announced an additional agreement with Moderna and the Government of Japan's Ministry of Health, Labour and Welfare (MHLW) to import and distribute an additional 50 million doses of COVID-19 Vaccine Moderna Intramuscular Injection in Japan from as early as the beginning of 2022. This agreement includes the potential to secure and supply vaccines corresponding to COVID-19 variants or booster products, should they be successfully developed by Moderna and licensed by the MHLW. Takeda will import and distribute the totaling 100 million doses including the additional 50 million doses in 2022 and 50 million doses announced in October, 2020.
- In July 2021, Takeda announced that the Japanese Ministry of Health, Labour and Welfare (MHLW) accepted the change in age indication in the package insert for COVID-19 Vaccine Moderna Intramuscular Injection to expand to 12 years of age and older. This change is based on the results of Moderna's Phase 2/3 study conducted in 3,732 subjects aged 12 to 17 years in the United States. The serum neutralizing antibody titer and neutralizing antibody titer response rate 28 days after the second vaccination of adolescents (12 to 17 years old), which are the primary endpoints, showed non-inferiority to young adults (18 to 25 years old) in the overseas phase 3 study (mRNA-1273-P301 study). Additionally, the results indicating a high preventive effect at the vaccine efficacy rate 2 weeks after the second vaccination, which was set as a secondary endpoint. No significant safety concerns were reported, as was the case with the results of clinical studies in patients aged 18 years or older.
- In December 2021, Takeda announced that Japan's Ministry of Health, Labour and Welfare (MHLW) has granted regulatory approval for a 50 µg booster dose of SPIKEVAX Intramuscular Injection, previously known as COVID-19 Vaccine Moderna Intramuscular Injection, in Japan for administration at least six months after completion of the primary series in those who are 18 years and older. The approval is based on previously-reported positive Moderna Phase 2 study results. Moderna's Phase 2 study was amended to offer a 50 µg booster dose to interested participants aged 18 years and older six to eight months following their second dose of the primary series of Moderna's COVID-19 vaccine. The results showed that a booster dose of the vaccine greatly increased neutralizing titers measured against the original virus strain compared to pre-boost levels. The reactogenicity profile observed following the booster dose was similar to the second dose of the primary series and the safety profile was also similar to that following any dose of Moderna's COVID-19 vaccine of the primary series.
- In December 2021, Takeda announced a third agreement with Japan's Ministry of Health, Labour and Welfare (MHLW) and Moderna to import and distribute 18 million additional doses of SPIKEVAX Intramuscular Injection in Japan in 2022. Takeda previously announced a three-way agreement with Moderna and MHLW to distribute 50 million doses of SPIKEVAX in Japan in 2021, and announced a second agreement for Takeda to import and distribute an additional 50 million doses in 2022, totaling 100 million doses between the two agreements. Due to the approval of the 50 microgram booster dose described in the foregoing paragraph, which is half of the dosage level used in the initial two-dose series of the vaccine (100 microgram per dose), the doses per vial for the second 50 million doses will increase, meaning Takeda will be able to deliver 75 million booster doses (at 15 doses per vial).

With this third agreement for 18 million doses (at 15 doses per vial), Takeda will now deliver a total of 93 million doses to Japan in 2022.

*Development code: NVX-CoV2373 (Japanese development code: TAK-019) / Generic name: COVID-19 vaccine*

- In September 2021, Takeda announced the agreement that the Japanese Ministry of Health, Labour and Welfare (MHLW) will purchase 150 million doses of Novavax, Inc. (Novavax)'s vaccine candidate (TAK-019 in Japan) manufactured in Japan by Takeda subject to licensing and approval. Takeda is establishing the capability to manufacture TAK-019 at its facilities in Japan and aims to begin distribution in early calendar year 2022. Novavax is licensing and transferring manufacturing technologies to enable Takeda to manufacture the vaccine antigen and is supplying the Matrix-M™ adjuvant to Takeda for fill/finish together with the antigen. Takeda is responsible for the Japanese clinical trial and regulatory submission and will distribute TAK-019 in Japan should it be approved by the MHLW.
- In December 2021, Takeda announced the submission of a New Drug Application (NDA) to the Ministry of Health, Labour and Welfare (MHLW) in Japan for TAK-019. The NDA submission includes an analysis from the ongoing Phase 1/2 immunogenicity and safety clinical trial of TAK-019 in Japan as well as safety and efficacy data from Novavax' global clinical trial program with more than 50,000 participants, including two pivotal Phase 3 trials, one conducted in the U.K. and another trial conducted in the U.S. and Mexico. Takeda's interim trial results showed that two 0.5 ml doses given 21 days apart induced robust anti-SARS-CoV-2 immune responses in healthy Japanese adults. No serious adverse events were reported in the TAK-019 group and the vaccine candidate was also well-tolerated. These results are consistent with previously reported global clinical trial results of Novavax' recombinant protein COVID-19 vaccine candidate (NVX-CoV2373). Takeda submitted all available Chemistry, Manufacturing and Controls (CMC), non-clinical and clinical data as of December 2021. Some additional CMC data will be subsequently submitted to the Japan Pharmaceuticals and Medical Devices Agency (PMDA) during the NDA review period. Through its partnership with Novavax including technology transfer, Takeda is establishing the capability to manufacture TAK-019 at its facilities in Japan and aims to begin distribution in early calendar year 2022, pending regulatory approval.

*Development code: TAK-003 / Generic name: Dengue vaccine*

- In May 2021, Takeda announced that TAK-003 demonstrated continued protection against dengue illness and hospitalization, regardless of an individual's previous dengue exposure, with no important safety risks identified through three years after vaccination in the ongoing pivotal Phase 3 Tetravalent Immunization against Dengue Efficacy Study (TIDES) trial. TIDES enrolled more than 20,000 healthy children and adolescents ages four to 16 years in dengue-endemic countries in Latin America and Asia. Safety and efficacy results from the 36-month follow-up exploratory analysis of TIDES were presented at the 17th Conference of the International Society of Travel Medicine (CISTM). Through three years (36 months after the second dose), observations of varied vaccine efficacy by serotype remained consistent with previously reported results. No evidence of disease enhancement was observed. TAK-003 was generally well tolerated, and there were no important safety risks observed. TIDES safety and efficacy data through 36-months follow-up was included in regulatory submissions to the European Union and dengue-endemic countries and will be part of additional filings planned for 2021, including in the United States.

***Building a sustainable research platform / Enhancing R&D collaboration***

In addition to our concentrated efforts to increase our in-house R&D capabilities, external partnerships with third-party partners are a key component of our strategy for enhancing our R&D pipeline. Our strategy to expand and diversify our external partnerships allows us to take part in research of a wide variety of new products and increases the chances that we will be able to take part in a major research-related breakthrough.

- In July 2021, Takeda and PeptiDream Inc. announced an expansion of its research collaboration and exclusive license agreement, announced in December 2020, to create peptide-drug conjugates (PDCs) for several central nervous system (CNS) targets, which play important roles in chronic neurodegenerative diseases. This new collaboration expands the use of the TfR1 binding peptide ligands for CNS targets associated with neurodegeneration allowing Takeda to conjugate the peptides with therapeutic cargoes optimized to cross the blood-brain barrier (BBB). A significant challenge to the development of effective medicines for neurodegenerative diseases is the ability to deliver therapeutic molecules across the BBB into the brain. Peptide carriers that bind to TfR1 when conjugated to various therapeutic payloads facilitate the transport of the payload across the BBB into the brain, and thereby significantly improve functional benefit. This TfR1 BBB shuttle approach has the potential to accelerate the development of therapies for which BBB penetration remains challenging. This approach may also enable broad brain region biodistribution that is frequently needed to effectively treat many neurodegenerative diseases for which few, if any, effective drugs currently exist.
- In July 2021, Takeda and Frazier Healthcare Partners announced a collaboration to launch HilleVax, Inc. (HilleVax), a biopharmaceutical company to develop and commercialize Takeda’s norovirus vaccine candidate. Takeda has granted a license to HilleVax for the exclusive development and commercialization rights to its norovirus vaccine candidate, HIL-214 (formerly TAK-214), worldwide outside of Japan, in exchange for upfront consideration, as well as future cash milestones and royalties on net sales. Takeda will retain commercialization rights in Japan and HilleVax will integrate certain Japan development activities into its global development. HIL-214, which is a virus-like particle (VLP) based vaccine candidate, completed a randomized, placebo-controlled Phase 2b field efficacy study in 4,712 adult subjects in which HIL-214 was well-tolerated and demonstrated clinical proof of concept in preventing moderate-to-severe cases of acute gastroenteritis from norovirus infection.<sup>1</sup> To date, the candidate has been studied in nine human clinical trials with safety data from over 4,500 subjects and immunogenicity data from over 2,000 subjects.
- In September 2021, Takeda and Mirum Pharmaceuticals, Inc. (Mirum) announced that the companies have entered into an exclusive licensing agreement for the development and commercialization of maralixibat chloride (maralixibat) (US trade name: LIVMARLI), an apical sodium dependent bile acid transporter (ASBT) inhibitor, in Japan for Alagille syndrome (ALGS), progressive familial intrahepatic cholestasis (PFIC), and biliary atresia (BA). Maralixibat, an investigational, orally administered medication, is being evaluated globally in ALGS, PFIC, and BA. Under the terms of the agreement, Takeda will be responsible for regulatory approval and commercialization of maralixibat in Japan. Takeda will also be responsible for development, including conducting clinical studies in cholestatic indications.
- In September 2021, Takeda and JCR Pharmaceuticals Co., Ltd. (JCR) announced a geographically-focused exclusive collaboration and license agreement to commercialize JR-141 (pabinafusp alfa), an investigational, next-generation recombinant fusion protein of an antibody against the human transferrin receptor and iduronate-2-sulfatase (IDS) enzyme for the treatment of Hunter syndrome (also known as Mucopolysaccharidosis type II or MPS II). JR-141, applied with J-Brain Cargo, JCR’s proprietary blood-brain barrier (BBB) technology, is engineered to transport the therapeutic enzyme across the BBB to directly reach the brain and address both the somatic and neuronopathic

manifestations of the disease, which can lead to progressive cognitive decline. Under the terms of the exclusive collaboration and license agreement, Takeda will exclusively commercialize JR-141 outside of the United States, including Canada, Europe, and other regions (excluding Japan and certain other Asia-Pacific countries). The two companies will collaborate to bring this therapy to patients as quickly as possible upon completion of the global Phase 3 program, which will be conducted by JCR. Takeda receives an option under a separate option agreement, which allows Takeda to acquire an exclusive license to commercialize JR-141 in the U.S. upon completion of the Phase 3 program.

- In October 2021, Takeda announced the exercise of its option to acquire GammaDelta Therapeutics Limited (“GammaDelta”), a company focused on exploiting the unique properties of gamma delta ( $\gamma\delta$ ) T cells for immunotherapy. Through the acquisition, Takeda will obtain GammaDelta’s allogeneic variable delta 1 (V $\delta$ 1) gamma-delta ( $\gamma\delta$ ) T cell therapy platforms, which includes both blood-derived and tissue-derived platforms, in addition to early-stage cell therapy programs. The deal is expected to be finalized in Q1 of Takeda’s fiscal year 2022. Closing of the transaction is contingent on completion of review under antitrust laws, including the Hart-Scott-Rodino Antitrust Improvements Act of 1976 (HSR) in the U.S. Takeda received HSR clearance for the transaction from the United States Federal Trade Commission (FTC) in January, 2022.
- In January 2022, Takeda announced the exercise of its option to acquire Adaptate Biotherapeutics Ltd. (“Adaptate”), a UK company focused on developing antibody-based therapeutics for the modulation of variable delta 1 (V $\delta$ 1) gamma delta ( $\gamma\delta$ ) T cells. Through the acquisition, Takeda will acquire Adaptate’s antibody-based  $\gamma\delta$  T cell engager platform, including pre-clinical candidate and discovery pipeline programs. Adaptate’s  $\gamma\delta$  T cell engagers are designed to specifically modulate  $\gamma\delta$  T cell-mediated immune responses at tumor sites while sparing damage to healthy cells. The planned acquisition of Adaptate follows Takeda’s recently exercised option to acquire GammaDelta Therapeutics and is intended to further accelerate the development of innovative  $\gamma\delta$  T cell-based therapies. The deal is expected to be finalized in Q1 of Takeda’s fiscal year 2022.

## Analysis of Results of Operations, Financial Position, and Cash Flow

### Results of Operations (Reported)

#### Consolidated Financial Results (April 1 to December 31, 2021)

	Billion JPY or percentage			
	FY2020 Q3YTD	FY2021 Q3YTD	Change versus the same period of the previous fiscal year	
Revenue	2,427.5	2,695.7	268.2	11.0 %
Cost of sales	(740.9)	(798.5)	(57.6)	7.8 %
Selling, general and administrative expenses	(641.3)	(662.9)	(21.7)	3.4 %
Research and development expenses	(342.5)	(382.5)	(39.9)	11.7 %
Amortization and impairment losses on intangible assets associated with products	(307.6)	(323.6)	(16.1)	5.2 %
Other operating income	118.5	34.3	(84.3)	(71.1)%
Other operating expenses	(155.1)	(100.0)	55.1	(35.5)%
Operating profit	358.7	462.5	103.7	28.9 %
Finance income and (expenses), net	(115.4)	(100.6)	14.8	(12.8)%
Share of loss of investments accounted for using the equity method	(8.0)	(5.3)	2.8	(34.4)%
Profit before tax	235.4	356.6	121.3	51.5 %
Income tax expenses	(56.3)	(115.1)	(58.7)	104.3 %
Net profit for the period	179.0	241.5	62.5	34.9 %

**Revenue.** Revenue for the nine-month period ended December 31, 2021 was 2,695.7 billion JPY, an increase of 268.2 billion JPY, or 11.0%, compared to the same period of the previous fiscal year. Excluding the impact from fluctuations in foreign exchange rates, which was calculated by translating revenue of the nine-month period ended December 31, 2021 using corresponding exchange rates in the same period of the previous fiscal year, the increase in revenue was 6.1%. In April 2021, Takeda completed the sale of a portfolio of diabetes products in Japan to Teijin Pharma Limited for 133.0 billion JPY, which was recorded as revenue and accounted for 5.5 percentage points (“pp”) of the increase in revenue. Excluding this selling price from revenue for the nine-month period ended December 31, 2021, the increase was 5.6%.

Each of our core therapeutic areas (i.e. Gastroenterology (“GI”), Rare Diseases, Plasma-Derived Therapies (“PDT”) Immunology, Oncology, and Neuroscience) contributed to positive revenue growth; however, Rare Diseases would have declined if not for the positive impact of the depreciation of the yen. Intensified competition impacted some products in this area, especially treatments for Rare Hematology. Overall, throughout the nine-month period ended December 31, 2021, the global spread of COVID-19 did not have a material effect on our revenue.

During the third quarter of the fiscal year ending March 31, 2022, LIVTENCITY (for post-transplant cytomegalovirus (“CMV”) infection/disease) was launched in the U.S. in December 2021, following the launch of EXKIVITY (for non-small cell lung cancer) in the U.S. in September 2021.

Revenue outside of our core therapeutic areas increased by 36.7 billion JPY, or 8.2%, compared to the same period of the previous fiscal year to 482.2 billion JPY, largely due to the 133.0 billion JPY selling price of the diabetes portfolio in Japan, offsetting the impact from divestitures. Revenue from distributing Moderna’s COVID-19 vaccine, SPIKEVAX Intramuscular Injection, in Japan also contributed to the growth.

Year-on-year change in revenue for this nine-month period in each of our main therapeutic areas was primarily attributable to the following products:

- *GI.* In Gastroenterology, revenue was 665.7 billion JPY, a year-on-year increase of 76.9 billion JPY, or 13.1%. Growth was driven by Takeda’s top-selling product ENTYVIO (for ulcerative colitis (“UC”) and Crohn’s disease (“CD”)), with sales of 395.4 billion JPY, a year-on-year increase of 76.1 billion JPY, or 23.8%. Sales in the U.S. increased by 46.7 billion JPY, or 21.3%, to 266.0 billion JPY driven by increase in the first line biologic inflammatory bowel disease (“IBD”) population both in UC and CD. Sales in Europe and Canada increased by 21.7 billion JPY, or 26.9%, to 102.2 billion JPY. In the Growth and Emerging Markets, the increase in sales was primarily driven by Brazil and China. Sales of TAKECAB (for acid-related diseases) were 78.4 billion JPY, an increase of 14.2 billion JPY, or 22.2%, versus the same period of the previous fiscal year. This increase was mainly driven by the expansion of new prescriptions in the Japanese market due to TAKECAB’s efficacy in reflux esophagitis and the prevention of recurrence of gastric and duodenal ulcers during low-dose aspirin administration. Sales of GATTEX/REVESTIVE (for short bowel syndrome) were 56.6 billion JPY, an



increase of 6.5 billion JPY, or 12.9%, primarily due to increased market penetration and new country launches including Japan. Sales of AMITIZA (for chronic constipation) decreased by 13.0 billion JPY, or 68.8%, to 5.9 billion JPY, due to generic entrants in the U.S. in January 2021.

- *Rare Diseases.* In Rare Diseases, revenue was 462.9 billion JPY, a year-on-year increase of 16.2 billion JPY, or 3.6%. Revenue in Rare Metabolic increased by 11.6 billion JPY, or 9.5%, compared to the same period of the previous fiscal year to 133.4 billion JPY. Sales of enzyme replacement therapies ELAPRASE (for Hunter syndrome), VPRIV (for Gaucher disease) and REPLAGAL (for Fabry disease) increased primarily in Europe and Growth and Emerging Markets. Revenue in Rare Hematology decreased by 7.0 billion JPY, or 3.2%, to 211.6 billion JPY. Sales of ADVATE decreased by 7.8 billion JPY, or 8.0%, to 89.3 billion JPY. Sales of ADYNOVATE/ADYNOVI increased by 2.1 billion JPY, or 4.8%, to 45.9 billion JPY, partially helped by the positive impact of the depreciation of the yen. Both products were impacted by the competitive landscape in the hemophilia A non-inhibitors market in the U.S. FEIBA sales decreased by 5.3 billion JPY, or 15.4%, to 29.0 billion JPY, negatively impacted by the difference in timing of government tenders in Growth and Emerging Markets. Revenue in Hereditary Angioedema (“HAE”) was 117.7 billion JPY, a year-on-year increase of 11.4 billion JPY, or 10.7%. Sales of TAKHZYRO were 78.4 billion JPY, an increase of 12.5 billion JPY, or 19.0%, versus the same period of the previous fiscal year primarily due to expansion of the prophylactic market, continued geographic expansion and strong patient uptake. Sales of CINRYZE decreased by 2.6 billion JPY, or 14.8%, to 14.7 billion JPY, primarily due to conversion to TAKHZYRO and a shift to newer agents marketed by competitors.
- *PDT Immunology.* In Plasma-Derived Therapies (“PDT”) Immunology, revenue increased by 50.2 billion JPY, or 16.0%, compared to the same period of the previous fiscal year to 363.2 billion JPY. Aggregate sales of immunoglobulin products were 278.3 billion JPY, an increase of 30.3 billion JPY, or 12.2%, compared to the same period of the previous fiscal year. In particular, sales of GAMMAGARD LIQUID (for the treatment of primary immunodeficiency (“PID”) and multifocal motor neuropathy (“MMN”)) increased due to higher demand versus the same period of the previous fiscal year. In addition, CUVITRU and HYQVIA, which are SCIG (subcutaneous immunoglobulin) therapies, marked double digit growth. Aggregate sales of albumin products including HUMAN ALBUMIN and FLEXBUMIN (primarily used for hypovolemia and hypoalbuminemia) were 61.5 billion JPY, an increase of 17.9 billion JPY, or 41.0%, versus the same period of the previous fiscal year driven by higher China sales following the resolution of the supply interruption which impacted HUMAN ALBUMIN for release in China in the second half of the previous fiscal year.
- *Oncology.* In Oncology, revenue was 359.1 billion JPY, a year-on-year increase of 40.6 billion JPY, or 12.8%. Sales of VELCADE (for multiple myeloma) increased by 8.6 billion JPY, or 11.3% versus the same period of the previous fiscal year to 84.5 billion JPY. U.S. sales increased by 9.6 billion JPY, or 13.3%, versus the same period of the previous fiscal year. This reflects a rebound in demand after lower sales in the first quarter of the previous fiscal year, when prescribers favored orally administered products over infusions or injections early in the COVID-19 pandemic. In addition, increased use of VELCADE as part of initial treatment for new patients contributed to the growth this year in the U.S. Royalty income outside the U.S. decreased due to continued generic erosion. Sales of NINLARO (for multiple myeloma) were 70.7 billion JPY, an increase of 2.9 billion JPY, or 4.2%, versus the same period of the previous fiscal year. In the U.S., NINLARO’s profile as an effective oral treatment led to a temporary increase in demand early in the COVID-19 pandemic in 2020 because its oral administration facilitated treatment in the at-home setting. This benefit has been less impactful in the U.S. this year; however, there have been strong demand increases in other countries, particularly in China. Sales of LEUPLIN/ENANTONE (generic name: leuprorelin) (for endometriosis, uterine fibroids, premenopausal breast cancer, prostatic cancer, etc.), an off-patented product, increased by 7.0 billion JPY, or 9.2%, versus the same period of the previous fiscal year to 82.2 billion JPY mainly driven by an increased supply in the U.S. which was partially offset by a decrease in Japan due to generic erosion and competition. Sales of ADCETRIS (for malignant lymphomas) increased by 7.4 billion JPY, or 16.7% versus the same period of the previous fiscal year to 51.8 billion JPY, led by strong growth in sales in Growth and Emerging Markets, particularly in China where it was approved in May 2020. Sales of ALUNBRIG (for non-small cell lung cancer) were 10.1 billion JPY, an increase of 3.6 billion JPY, or 56.2% due to new launches and market penetration around the world.
- *Neuroscience.* In Neuroscience, revenue was 362.6 billion JPY, a year-on-year increase of 47.5 billion JPY, or 15.1%. Sales of VYVANSE/ELVANSE (for attention deficit hyperactivity disorder (“ADHD”)) were 245.0 billion JPY, an increase of 42.6 billion JPY, or 21.0%, versus the same period of the previous fiscal year. VYVANSE/ELVANSE has been negatively affected by COVID-19 during the course of the pandemic, most notably during periods when stay-at-home restrictions have been in place reducing patient visits, subsequent diagnoses and creating temporary discontinuation of medication. The trend has been fluctuating throughout 2020 and into 2021; however, there has been a positive impact from increasing prescriptions versus the same period of the previous fiscal year. Sales of TRINTELLIX (for major depressive disorder (“MDD”)) were 63.0 billion JPY, an increase of 10.4 billion JPY, or 19.6%, versus the same period of the

previous fiscal year, primarily due to increasing prescriptions in the U.S. and in Japan. The increase of these products was partially offset by the decrease of other neuroscience products such as REMINYL (for Alzheimer's disease), attributable to the continued impact of competition from generic products.

Revenue by Geographic Region:

Revenue:	Billion JPY; percentages are portion of total revenue			
	FY2020 Q3YTD		FY2021 Q3YTD	
Japan <sup>*1</sup>	435.1	17.9 %	530.2	19.7 %
United States	1,189.0	49.0 %	1,297.0	48.1 %
Europe and Canada	500.0	20.6 %	541.0	20.1 %
Asia (excluding Japan)	119.2	4.9 %	139.8	5.2 %
Latin America	95.4	3.9 %	93.5	3.5 %
Russia/CIS	38.7	1.6 %	43.6	1.6 %
Other <sup>*2</sup>	50.2	2.1 %	50.6	1.9 %
Total	2,427.5	100.0 %	2,695.7	100.0 %

\*1 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in the nine-month period ended December 31, 2021.

\*2 Other includes the Middle East, Oceania and Africa.

**Cost of Sales.** Cost of Sales increased by 57.6 billion JPY, or 7.8%, to 798.5 billion JPY. The increase was primarily due to the depreciation of the yen and a sales increase of products with higher cost of sales ratio as compared to the same period of the previous fiscal year. The increase was partially offset by a 42.5 billion JPY decrease in non-cash charges related to the unwind of the fair value step up on acquired inventory recognized in connection with the acquisition of Shire plc. The Cost of Sales Ratio decreased by 0.9 pp compared to the same period of the previous fiscal year to 29.6%. The main reason for the decrease in the Cost of Sales Ratio was the effect of the sale of a portfolio of diabetes products in Japan with the selling price of 133.0 billion JPY being recorded in revenue.

**Selling, General and Administrative (SG&A) expenses.** SG&A expenses increased by 21.7 billion JPY, or 3.4%, to 662.9 billion JPY compared to the same period of the previous fiscal year, mainly due to the impact from the depreciation of the yen in the current period.

**Research and Development (R&D) expenses.** R&D expenses increased by 39.9 billion JPY, or 11.7%, to 382.5 billion JPY compared to the same period of the previous fiscal year, mainly due to further investment in prioritized new molecular entities as well as the impact from the depreciation of the yen in the current period.

**Amortization and Impairment Losses on Intangible Assets Associated with Products.** Amortization and Impairment Losses on Intangible Assets Associated with Products increased by 16.1 billion JPY, or 5.2%, to 323.6 billion JPY compared to the same period of the previous fiscal year mainly due to impairment charges related to certain in-process R&D assets recorded in the current period.

**Other Operating Income.** Other Operating Income was 34.3 billion JPY, a decrease of 84.3 billion JPY, or 71.1%, compared to the same period of the previous fiscal year, mainly driven by a 60.2 billion JPY revaluation gain recorded in the same period of the previous fiscal year triggered by an update to previously recognized liabilities for pipeline compound SHP647 and certain associated rights ("SHP647"), to reflect management's decision to terminate the clinical trial program following the European Commission's decision in May 2020 to release Takeda's obligation to divest SHP647. The decrease was also driven by the effect of a 37.2 billion JPY divestiture gain on the sale of non-core assets in Asia Pacific, Europe, and Canada recorded in the same period of the previous fiscal year.

**Other Operating Expenses.** Other Operating Expenses were 100.0 billion JPY, a decrease of 55.1 billion JPY, or 35.5%, compared to the same period of the previous fiscal year. This is mainly attributable to a 27.3 billion JPY decrease in restructuring expenses mainly attributable to lower Shire integration costs. There was also a 18.7 billion JPY loss recognized in the same period of the previous year from changes in the fair value of financial assets associated with contingent consideration arrangements from the divestment of XIIDRA.

**Operating Profit.** As a result of the above factors, Operating Profit increased by 103.7 billion JPY, or 28.9% compared to the same period of the previous fiscal year to 462.5 billion JPY.

**Net Finance Expenses.** Net Finance Expenses were 100.6 billion JPY in the current period, a decrease of 14.8 billion JPY, or 12.8%, compared to the same period of the previous fiscal year. The decrease is mainly due to a decrease in interest expense primarily driven by the reduction in outstanding balances of bonds and loans and a gain on prior equity method investments related to the acquisition of Maverick Therapeutics, Inc. in April 2021.

**Share of Loss of Investments Accounted for Using the Equity Method.** Share of Loss of Investments Accounted for Using the Equity Method was 5.3 billion JPY, a decrease of 2.8 billion JPY, or 34.4%, compared to the same period of the previous fiscal year, mainly due to Takeda's share of impairment loss recognized by Teva Takeda Pharma Ltd. in the same period of the previous fiscal year resulting from the reassessment of the recoverable amount of relevant assets triggered by the decision to divest a part of its generics business and a manufacturing plant.

**Income Tax Expenses.** Income Tax Expenses were 115.1 billion JPY, an increase of 58.7 billion JPY, or 104.3%, compared to the same period of the previous year. This increase was primarily due to a tax charge of 64.6 billion JPY for tax and interest, net of 0.5 billion JPY of associated tax benefit, arising from tax assessment involving Irish taxation of the break fee Shire received from AbbVie in connection with the terminated offer to acquire Shire made by AbbVie in 2014 as well as higher pretax earnings in the current period. These increases were partially offset by an increase in tax credits and a decrease in unitary tax in overseas subsidiaries in the current period versus the same period of the previous year.

**Net Profit for the Period.** Net Profit for the Period increased by 62.5 billion JPY, or 34.9%, compared to the same period of the previous fiscal year to 241.5 billion JPY.



## **Results of Operations (Underlying) (April 1 to December 31, 2021)**

### ***Definition of Core and Underlying Growth***

Takeda uses the concept of Underlying Growth for internal planning and performance evaluation purposes.

Underlying Growth compares two periods (fiscal quarters or years) of financial results under a common basis and is used by management to assess the business. These financial results are calculated on a constant currency basis using a full year plan rate and exclude the impacts of divestitures and other amounts that are unusual, non-recurring items or unrelated to our ongoing operations. Although these are not measures defined by IFRS, Takeda believes Underlying Growth is useful to investors as it provides a consistent measure of our performance.

Takeda uses "Underlying Revenue Growth", "Underlying Core Operating Profit Growth", and "Underlying Core EPS Growth" as key financial metrics.

Underlying Revenue represents revenue on a constant currency basis and excluding non-recurring items and the impact of divestitures that occurred during the reported periods presented.

Underlying Core Operating Profit represents Core Operating Profit (as defined below) on a constant currency basis and further adjusted to exclude the impacts of divestitures that occurred during the reporting periods presented.

Underlying Core EPS represents net profit based on a constant currency basis, adjusted to exclude the impact of divestitures, items excluded in the calculation of Core EPS (as defined below), divided by the outstanding shares (excluding treasury shares) as of the end of the comparative period.

Core Revenue represents revenue adjusted to exclude significant items unrelated to Takeda's core operations.

Core Operating Profit represents net profit adjusted to exclude income tax expenses, the share of profit or loss of investments accounted for using the equity method, finance expenses and income, other operating expenses and income, amortization and impairment losses on acquired intangible assets and other items unrelated to Takeda's core operations, such as non-recurring items, purchase accounting effects and transaction related costs.

Core EPS represents net profit adjusted to exclude the impact of items excluded in the calculation of Core Operating Profit, and other non-operating items (e.g. amongst other items, fair value adjustments and the imputed financial charge related to contingent consideration) that are unusual, non-recurring in nature or unrelated to Takeda's ongoing operations and the tax effect of each of the adjustments, divided by the average outstanding shares (excluding treasury shares) of the reporting periods presented.

### ***Underlying Results***

#### **FY2021 Q3YTD**

Underlying Revenue Growth	+7.1%
Underlying Core Operating Profit Growth	+5.4%
Underlying Core Operating Profit Margin	29.4%
Underlying Core EPS Growth	+9.9%

***Underlying Revenue Growth*** was 7.1% compared to the same nine-month period of the previous fiscal year. Underlying revenue attributable to Takeda's 14 global brands\* grew by 12.0%, which constitute approximately 42% of the total Underlying revenue, led by ENTYVIO, TAKHZYRO, HUMAN ALBUMIN/FLEXBUMIN, GAMMAGARD LIQUID/KIOVIG and CUVITRU.

\* Takeda's 14 global brands

GI: ENTYVIO, GATTEX/REVESTIVE, ALOFISEL

Rare Diseases: NATPARA/NATPAR, ADYNOVATE/ADYNOVI, TAKHZYRO, ELAPRASE, VPRIV

PDT Immunology: GAMMAGARD LIQUID/KIOVIG, HYQVIA, CUVITRU, HUMAN ALUBUMIN/FLEXBUMIN

Oncology: NINLARO, ALUNBRIG

**Underlying Revenue Growth by Therapeutic Area**

GI	+7.6%
Rare Diseases	-1.0%
Rare Metabolic	+5.2%
Rare Hematology	-7.6%
Hereditary Angioedema	+5.4%
PDT Immunology	+10.3%
Oncology	+8.2%
Neuroscience	+10.0%
Other	+10.6%
Total	+7.1%

(Note) Underlying Revenue represents revenue on a constant currency basis and excluding non-recurring items and the impact of divestitures. Please refer to "[Results of Operations \(Reported\)](#)" for the revenue of each core therapeutic areas and sales of major products before underlying adjustments.

The impact of major non-recurring items and divestitures excluded to calculate Underlying Revenue:

- Revenue of select over-the-counter and non-core products in Asia Pacific is excluded from the same period of the previous fiscal year as the divestiture was completed in November 2020.
- Revenue of select non-core prescription pharmaceutical products predominantly in Europe is excluded from the same period of the previous fiscal year as the divestiture was completed in December 2020.
- Revenue of select over-the-counter and non-core products in Latin America is excluded from the same period of the previous fiscal year as the divestiture was completed in January 2021.
- Net sales from TACHOSIL, a surgical patch, are excluded from the same period of the previous fiscal year as the divestiture was completed in January 2021.
- Revenue of select over-the-counter and non-core products predominantly in Europe is excluded from the same period of the previous fiscal year as the divestiture was completed in March 2021.
- Revenue of the former subsidiary, Takeda Consumer Healthcare Company Limited is excluded from the same period of the previous fiscal year as the divestiture was completed in March 2021.
- Net sales from a portfolio of diabetes products in Japan (NESINA, LIOVEL, INISYNC and ZAFATEK) are excluded from the same period of the previous fiscal year as the divestiture was completed at the beginning of April 2021. In addition, the non-recurring item of the 133.0 billion JPY selling price as the result of the completion of the divestiture is excluded from the current period.
- Revenue of select non-core prescription pharmaceutical products in China is excluded from both the current period and the same period of the previous fiscal year as the divestiture was publicly announced and had been expected to complete within the first half of the current fiscal year. It is now expected to complete in the second half of the current fiscal year.

**Underlying Core Operating Profit Growth** was 5.4% over the same nine-month period of the previous fiscal year, attributable to Underlying Revenue Growth.

Core Operating Profit for the current period, which excludes items unrelated to Takeda's core operations such as the sale of a portfolio of diabetes products in Japan, was 757.9 billion JPY.

**Underlying Core Operating Profit Margin** for the current period was 29.4%.

**Underlying Core EPS Growth** for the current period was 9.9%.

## **Consolidated Financial Position**

**Assets.** Total Assets as of December 31, 2021 were 12,698.5 billion JPY, reflecting a decrease of 213.8 billion JPY compared to the previous fiscal year-end. Cash and cash equivalents decreased by 241.9 billion JPY and Intangible Assets decreased by 117.2 billion JPY mainly due to amortization. These decreases were partially offset by an increase in Goodwill of 134.1 billion JPY mainly due to the effect of foreign currency translation.

Although there was a decline in share price during the three-month period ended December 31, 2021 that eliminated our surplus in market capitalization compared to the carrying value of our one cash-generating unit (CGU), we concluded there was no indication of goodwill impairment as of December 31, 2021.

**Liabilities.** Total Liabilities as of December 31, 2021 were 7,366.7 billion JPY, reflecting a decrease of 368.4 billion JPY compared to the previous fiscal year-end. Bonds and Loans decreased by 280.5 billion JPY to 4,354.9 billion JPY\* primarily as a result of the repayment of loans and the redemption of bonds. In addition, Other Financial Liabilities decreased by 114.7 billion JPY.

\* The carrying amount of Bonds was 3,653.7 billion JPY and Loans was 701.2 billion JPY as of December 31, 2021. Breakdown of Bonds and Loans carrying amount is as follows.

### Bonds:

<b>Name of Bond (Face Value if Denominated in Foreign Currency)</b>	<b>Issuance</b>	<b>Maturity</b>	<b>Carrying Amount (Billion JPY)</b>
Unsecured US dollar denominated senior notes (1,520 million USD)	June 2015	June 2022 ~ June 2045	175.1
Unsecured US dollar denominated senior notes (5,500 million USD)	September 2016	September 2023 ~ September 2026	606.6
Unsecured Euro denominated senior notes (3,750 million EUR)	November 2018	November 2022 ~ November 2030	486.3
Unsecured US dollar denominated senior notes (3,250 million USD)	November 2018	November 2023 ~ November 2028	372.3
Hybrid bonds (subordinated bonds)	June 2019	June 2079	498.0
Unsecured US dollar denominated senior notes (7,000 million USD)	July 2020	March 2030 ~ July 2060	800.1
Unsecured Euro denominated senior notes (3,600 million EUR)	July 2020	July 2027 ~ July 2040	466.0
Unsecured JPY denominated senior bonds	October 2021	October 2031	249.3
<b>Total</b>			<b>3,653.7</b>

Loans:

Name of Loan (Face Value if Denominated in Foreign Currency)	Execution	Maturity	Carrying Amount (Billion JPY)
Syndicated loans	April 2016	April 2023 ~ April 2026	200.0
Syndicated loans	April 2017	April 2027	113.5
Syndicated loans (1,500 million USD)	April 2017	April 2027	172.4
Bilateral loans	March 2016 ~ April 2017	March 2023 ~ March 2026	210.0
Other			5.3
Total			701.2

On May 17, 2021, Takeda redeemed the remaining 200 million USD of unsecured U.S. dollar-denominated senior notes issued in July 2017 in advance of their original maturity date of January 18, 2022. Following this, on June 11, 2021, Takeda prepaid 2,000 million USD of the Japan Bank for International Cooperation loan (“JBIC Loan”) amount of 3,700 million USD (that was entered into on December 3, 2018) in advance of its original maturity date of December 11, 2025. On August 10, 2021, Takeda redeemed 1,500 million EUR of unsecured senior notes issued in November 2018 in advance of their original maturity date of November 21, 2022. On October 14, 2021, Takeda issued 10-year unsecured senior bonds with an aggregate principal amount of 250 billion JPY and a maturity date of October 14, 2031. Following this, on December 13, 2021 Takeda prepaid the remaining 1,700 million USD amount outstanding on the JBIC Loan in advance of its original maturity date of December 11, 2025.

**Equity.** Total Equity as of December 31, 2021 was 5,331.8 billion JPY, an increase of 154.6 billion JPY compared to the previous fiscal year-end. This was mainly due to an increase of 215.5 billion JPY in Other Components of Equity mainly due to fluctuation in currency translation adjustments reflecting the depreciation of yen. This increase was partially offset by a decrease in Retained Earnings of 43.0 billion JPY and an increase in Treasury Shares of 31.5 billion JPY. The decrease in Retained Earnings resulted primarily from dividend payments of 284.2 billion JPY whereas Net Profit for the Period was recorded.

## **Consolidated Cash Flow**

	Billion JPY	
	FY2020 Q3YTD	FY2021 Q3YTD
Net cash from (used in) operating activities	610.0	747.5
Net cash from (used in) investing activities	100.2	(172.5)
Net cash from (used in) financing activities	(718.3)	(826.5)
Net increase (decrease) in cash and cash equivalents	(8.1)	(251.4)
Cash and cash equivalents at the beginning of the year	637.6	966.2
Effects of exchange rate changes on cash and cash equivalents	(11.8)	9.5
Net increase (decrease) in cash and cash equivalents resulting from a transfer from (to) assets held for sale	(0.1)	—
Cash and cash equivalents at the end of the period	617.6	724.3

*Net cash from operating activities* was 747.5 billion JPY for the current period compared to 610.0 billion JPY for the same period of the previous year. The increase of 137.6 billion JPY was primarily driven by higher net profit for the period adjusted for non-cash items and other adjustments, including the income relating to the release from the obligation to divest the pipeline compound SHP 647 and certain associated rights in the same period of the previous year. In addition, there was a decrease in trade and other receivables mainly due to the trade receivables sales program put in place in the current period. These favorable impacts were partially offset by a decrease in provisions due to payments.

*Net cash used in investing activities* was 172.5 billion JPY for the current period compared to net cash from investing activities of 100.2 billion JPY for the same period of the previous year. This increase in net cash used of 272.7 billion JPY was mainly due to a decrease of 122.8 billion JPY in proceeds from sales of business, net of cash and cash equivalents divested reflecting the sales of the non-core assets in the same period of the previous year, a decrease of 57.7 billion JPY in proceeds from sales and redemptions of investments, and an increase of 49.7 billion JPY in acquisition of businesses, net of cash and cash equivalents acquired.

*Net cash used in financing activities* was 826.5 billion JPY for the current period compared to 718.3 billion JPY for the same period of the previous year. The increase of 108.2 billion JPY was mainly due to a decrease in proceeds from issuance of bonds and long-term loans of 930.2 billion JPY as a result of the issuance of U.S. dollar-denominated senior notes of 7,000 million USD and Euro-denominated senior notes of 3,600 million EUR in the same period of the previous year compared to the 250 billion JPY issuance of senior bond in the current period. In addition, purchase of treasury shares increased by 50.4 billion JPY mainly due to the share buybacks in the current period. These were partially offset by a decrease in repayments of bonds and long-term loans of 754.1 billion JPY and the favorable impact from short-term loans and commercial papers of 85.0 billion JPY.

## Outlook for the Fiscal Year Ending March 31, 2022

Considering Takeda's financial results through the nine-month period ended December 31, 2021, the full-year financial forecast for revenue and profits have been revised from the previous forecast announced on October 28, 2021, reflecting favorable foreign exchange rates, strong business momentum driven by Takeda's 14 global brands, and discipline in operating expenses. The forecast upgrade is also attributable to other assumptions including updated tax rate assumptions.

### Full Year Reported Forecast for the Fiscal Year Ending March 31, 2022 (FY2021)

	Billion JPY or percentage			
	Previous Forecast (October 28, 2021)	Revised Forecast (February 3, 2022)	vs. Previous Forecast	
Revenue	3,370.0	3,510.0	+140.0	+4.2 %
Operating profit	488.0	515.0	+27.0	+5.5 %
Profit before tax	352.0	385.0	+33.0	+9.4 %
Net profit for the year (attributable to owners of the Company)	184.3	242.5	+58.2	+31.6 %
EPS (JPY)	117.35	154.99	+37.64	+32.1 %
Core Operating Profit	930.0	970.0	+40.0	+4.3 %
Core EPS (JPY)	394	416	+22	+5.6 %

### Major assumptions used in preparing the FY2021 Revised Reported Forecast

Major assumptions used in preparing the FY2021 reported forecast have been revised as below, including free cash flow, reflecting a strong outlook for the year.

	Billion JPY or percentage	
	Previous Forecast (October 28, 2021)	Revised Forecast (February 3, 2022)
FX rates	1 USD = 108 JPY 1 Euro = 131 JPY 1 RUB = 1.4 JPY 1 BRL = 19.9 JPY 1 CNY = 16.8 JPY	1 USD = 111 JPY 1 Euro = 131 JPY 1 RUB = 1.5 JPY 1 BRL = 20.7 JPY 1 CNY = 17.3 JPY
R&D expenses	(522.0)	(522.0)
Amortization of intangible assets associated with products	(406.0)	(412.0)
Of which Shire acquisition related	(328.0)	(335.0)
Impairment of intangible assets associated with products	(50.0)	(40.0)
Other operating income	23.0	48.0
Other operating expenses	(100.0)	(150.0)
Japan diabetes portfolio divestiture gain	130.0	131.4
Other Core Operating Profit adjustments	(39.0)	(32.4)
Of which Shire acquisition related to unwind of inventories step-up	(31.1)	(31.8)
Finance income and (expenses), net	(130.0)	(121.0)
Free cash flow (including announced divestitures)	600.0 - 700.0	700.0 - 800.0
Capital expenditures (cash flow base)	(210.0 - 260.0)	(210.0 - 260.0)
Depreciation and amortization (excluding intangible assets associated with products)	(150.0)	(150.0)
Cash tax rate on adjusted EBITDA (excluding divestitures)	Mid-teen%	Mid-teen%

**Management Guidance\***

Management guidance has not been changed from the previous guidance.

	<b>Guidance as of October 28, 2021</b>	<b>Guidance as of February 3, 2022</b>
Underlying Revenue Growth	Mid-single-digit growth	Mid-single-digit growth
Underlying Core Operating Profit Growth	Mid-single-digit growth	Mid-single-digit growth
Underlying Core Operating Profit Margin	~30% margin	~30% margin
Underlying Core EPS Growth	Mid-single-digit growth	Mid-single-digit growth

\* Please refer to "[Results of Operations \(Underlying\) \(April 1 to December 31, 2021\)](#)", Definition of Core and Underlying Growth.

**Other assumptions used in preparing the FY2021 Reported Forecast and the Management Guidance**

- To date, Takeda has not experienced a material effect on its financial results as a result of the global spread of the novel coronavirus infectious disease (COVID-19). Based on currently available information, Takeda believes that its financial results for FY2021 will not be materially affected by COVID-19 and, accordingly, Takeda's FY2021 forecast reflects this belief. However, the situation surrounding COVID-19 remains highly fluid, and future COVID-19-related developments in FY2021, including new or additional COVID-19 outbreaks and additional or extended lockdowns, shelter-in-place orders or other government action in major markets, could result in further or more serious disruptions to Takeda's business, such as slowdowns in demand for Takeda's products, supply chain related issues or significant delays in its clinical trial programs. These events, if they occur, could result in an additional impact on Takeda's business, results of operations or financial condition, as well as result in significant deviations from Takeda's FY2021 forecast.
- Takeda expects at least one 505(b)2 competitor for subcutaneous VELCADE to launch in the U.S. in late FY2021.
- Takeda does not expect to restart sales of NATPARA in the U.S. market in FY2021.
- The forecast and the guidance do not include the impact of any potential further divestitures beyond what has already been disclosed by Takeda.

**Forward looking statements**

All forecasts in this document are based on information currently available to management, and do not represent a promise or guarantee to achieve these forecasts. Various uncertain factors could cause actual results to differ, such as changes in the business environment and fluctuations in foreign exchange rates. Should any significant event occur which requires the forecast to be revised, the Company will disclose it in a timely manner.



## **Impact of the Spread of the Novel Coronavirus Infectious Disease (COVID-19) and Takeda's Initiatives in Response**

### **(i) Impact of COVID-19 on Takeda's Operations and Financial Condition**

Takeda continues to respond to the COVID-19 pandemic and provide industry support in a number of ways. While vaccines are becoming more broadly available, we continue to strictly adhere to local public health guidance across our geographies in addition to the existing protocols we have had in place for almost two years, and monitor any potential impacts of effects and evolution of COVID-19, including new variants like Omicron, on our business activities.

In monitoring demand for our products, we have seen limited impact as many of our medicines are for severe chronic or life-threatening diseases, without the requirement of a hospital elective procedure. In terms of our global supply chain, based on current assessments, we have not seen, nor do we anticipate, any material potential supply distribution issues due to the COVID-19 outbreak. Where appropriate and in accordance with local public health guidance and regulations, our field employees have resumed some face-to-face engagements with customers, with the majority of all interactions still virtual. Clinical trial activities that were temporarily paused during the previous fiscal year have generally been resumed while we continue to monitor the evolution of the pandemic.

As we continue to monitor developments in the financial markets, we currently do not anticipate any material liquidity or funding-related issues.

### **(ii) Takeda's Initiatives to Mitigate the Impact of COVID-19**

Guided by our values, Takeda's response to COVID-19 continues to focus on protecting the health and safety of our employees, our ability to ensure our medicines are available to patients who rely on them and playing our part to reduce transmission and support the communities where our employees live and work.

Major updates to Takeda's initiatives in response to the spread of COVID-19 in the current period are as below.

- The highly contagious Omicron variant has temporarily slowed the roll out of a new hybrid working model in parts of the business. Moving forward, implementation of this model will vary by job function, and on the local level, given differences in public health guidance and regulations, changes in population and epidemiology over time and standards of practice in the community.
- Takeda has undertaken a number of efforts to help the world respond to COVID-19. This includes bringing COVID-19 vaccines to Japan through two partnerships. The first partnership is with Novavax, for the development, manufacturing, and commercialization of its COVID-19 vaccine candidate, NVX-CoV2373 (development code in Japan: TAK-019) in Japan. In September 2021, Takeda concluded the agreement with the Government of Japan's Ministry of Health, Labour and Welfare (MHLW) to provide 150 million doses of Novavax' COVID-19 vaccine candidate manufactured in Japan by Takeda subject to licensing and approval, starting in early calendar year 2022. In December 2021, Takeda submitted a New Drug Application (NDA) to the MHLW in Japan for TAK-019.

The second partnership is with Moderna and the MHLW to import and distribute Moderna's mRNA COVID-19 vaccine (SPIKEVAX Intramuscular Injection (former product name: COVID-19 Vaccine Moderna Intramuscular Injection)) in Japan. Since May 2021, Takeda has been distributing the Moderna COVID-19 vaccine in Japan. In October 2021, Takeda and Moderna published an investigation report prompted by the recall of three lots of the Moderna COVID-19 vaccine in Japan based on the observation of foreign particles in unpunctured vials from a single lot. The report concluded that the event does not pose an undue risk to patient safety or adversely affect the benefit/risk profile of the product.

Takeda will continue distribution of the vaccine in Japan in 2022 via an additional three-way agreement with Moderna and the MHLW. Specifically, the parties reached to an agreement in December 2021, to import and distribute 18 million additional doses of Moderna's COVID-19 vaccine, bringing the total to 93 million doses in 2022.

### **(iii) FY2021 Q3YTD financial impact from COVID-19**

Overall, the global spread of COVID-19 did not have a material effect on our financials for the nine-month period ended December 31, 2021. Over the course of the pandemic, there have been adverse effects due to COVID-19 observed in certain therapeutic areas, especially in Neuroscience during periods when stay-at-home restrictions have been in place, reducing patient visits to medical care providers. This was notable especially in the early months of the previous fiscal year. The trend has fluctuated since then, and we have not yet seen a full recovery to pre-COVID-19 levels, however, a certain number of our life-saving medicines have shown resilience and have grown even under such an environment. We have not experienced a material disruption from the rapid spread of COVID-19 due to the Omicron variant.



## Consolidated Financial Statements [IFRS]

### (1) Consolidated Statements of Profit or Loss

	JPY (millions, except per share data)		USD (millions) <sup>(*)</sup>
	Nine-month Period Ended December 31,		Nine-month Period Ended December 31,
	2020	2021	2021
Revenue	¥ 2,427,538	¥ 2,695,717	\$ 23,406
Cost of sales	(740,862)	(798,466)	(6,933)
Selling, general and administrative expenses	(641,275)	(662,932)	(5,756)
Research and development expenses	(342,544)	(382,459)	(3,321)
Amortization and impairment losses on intangible assets associated with products	(307,570)	(323,632)	(2,810)
Other operating income	118,532	34,269	298
Other operating expenses	(155,090)	(100,034)	(869)
Operating profit	358,729	462,463	4,015
Finance income	58,030	42,949	373
Finance expenses	(173,389)	(143,539)	(1,246)
Share of loss of investments accounted for using the equity method	(8,013)	(5,255)	(46)
Profit before tax	235,357	356,618	3,096
Income tax expenses	(56,330)	(115,077)	(999)
Net profit for the period	179,027	241,541	2,097
Attributable to:			
Owners of the Company	178,907	241,417	2,096
Non-controlling interests	120	124	1
Net profit for the period	179,027	241,541	2,097
Earnings per share (JPY)			
Basic earnings per share	114.57	154.09	1.34
Diluted earnings per share	113.72	153.03	1.33

(\*) Consolidated statements of profit or loss have been translated solely for the convenience of the reader at an exchange rate of 1USD = 115.17 JPY, the Noon Buying Rate certified by the Federal Reserve Bank of New York on December 30, 2021. The rate and methodologies used for the convenience translations differ from the currency exchange rates and translation methodologies under IFRS used for the preparation of the interim consolidated financial statements. The translation should not be construed as a representation that the Japanese yen amounts could be converted into U.S. dollars at the above or any other rate.

## (2) Consolidated Statements of Comprehensive Income

	JPY (millions)		USD (millions) <sup>(*)</sup>
	Nine-month Period Ended December 31,		Nine-month Period Ended December 31,
	2020	2021	2021
Net profit for the period	¥ 179,027	¥ 241,541	\$ 2,097
Other comprehensive income (loss)			
Items that will not be reclassified to profit or loss:			
Changes in fair value of financial assets measured at fair value through other comprehensive income	69,336	(5,951)	(52)
Remeasurement of defined benefit pension plans	(4,879)	(2,912)	(25)
	64,457	(8,862)	(77)
Items that may be reclassified subsequently to profit or loss:			
Exchange differences on translation of foreign operations	(42,370)	206,582	1,794
Cash flow hedges	(21,596)	13,958	121
Hedging cost	(10,288)	5,969	52
Share of other comprehensive income (loss) of investments accounted for using the equity method	220	(145)	(1)
	(74,034)	226,365	1,965
Other comprehensive income (loss) for the period, net of tax	(9,577)	217,503	1,889
Total comprehensive income for the period	169,450	459,044	3,986
Attributable to:			
Owners of the Company	169,301	458,887	3,984
Non-controlling interests	149	157	1
Total comprehensive income for the period	169,450	459,044	3,986

(\*) Consolidated statements of comprehensive income have been translated solely for the convenience of the reader at an exchange rate of 1USD = 115.17 JPY, the Noon Buying Rate certified by the Federal Reserve Bank of New York on December 30, 2021. The rate and methodologies used for the convenience translations differ from the currency exchange rates and translation methodologies under IFRS used for the preparation of the interim consolidated financial statements. The translation should not be construed as a representation that the Japanese yen amounts could be converted into U.S. dollars at the above or any other rate.

### (3) Consolidated Statements of Financial Position

	JPY (millions)		USD (millions) <sup>(*)</sup>
	As of March 31, 2021	As of December 31, 2021	As of December 31, 2021
<b>ASSETS</b>			
Non-current assets:			
Property, plant and equipment	¥ 1,453,917	¥ 1,493,587	\$ 12,969
Goodwill	4,033,917	4,167,993	36,190
Intangible assets	3,909,106	3,791,875	32,924
Investments accounted for using the equity method	112,468	104,507	907
Other financial assets	235,882	230,305	2,000
Other non-current assets	100,341	79,645	692
Deferred tax assets	353,769	352,715	3,063
Total non-current assets	10,199,400	10,220,626	88,744
Current assets:			
Inventories	753,881	811,324	7,045
Trade and other receivables	783,091	715,515	6,213
Other financial assets	36,598	27,555	239
Income taxes receivable	29,623	40,602	353
Other current assets	122,789	138,352	1,201
Cash and cash equivalents	966,222	724,341	6,289
Assets held for sale	20,689	20,203	175
Total current assets	2,712,893	2,477,893	21,515
Total assets	12,912,293	12,698,519	110,259
<b>LIABILITIES AND EQUITY</b>			
<b>LIABILITIES</b>			
Non-current liabilities:			
Bonds and loans	4,613,218	4,231,939	36,745
Other financial liabilities	517,677	461,692	4,009
Net defined benefit liabilities	158,857	169,803	1,474
Income taxes payable	33,690	30,874	268
Provisions	38,748	34,042	296
Other non-current liabilities	56,898	70,486	612
Deferred tax liabilities	542,852	558,607	4,850
Total non-current liabilities	5,961,940	5,557,443	48,254
Current liabilities:			
Bonds and loans	22,153	122,936	1,067
Trade and other payables	343,838	351,185	3,049
Other financial liabilities	248,053	189,298	1,644
Income taxes payable	145,203	185,441	1,610
Provisions	471,278	421,481	3,660
Other current liabilities	542,651	538,913	4,679
Total current liabilities	1,773,176	1,809,254	15,709
Total liabilities	7,735,116	7,366,697	63,964

	JPY (millions)		USD (millions) <sup>(*)</sup>
	As of March 31, 2021	As of December 31, 2021	As of December 31, 2021
<b><u>EQUITY</u></b>			
Share capital	1,668,145	1,676,263	14,555
Share premium	1,688,424	1,697,562	14,740
Treasury shares	(59,552)	(91,013)	(790)
Retained earnings	1,509,906	1,466,926	12,737
Other components of equity	366,114	581,592	5,050
Equity attributable to owners of the company	5,173,037	5,331,330	46,291
Non-controlling interests	4,140	493	4
Total equity	5,177,177	5,331,822	46,295
Total liabilities and equity	12,912,293	12,698,519	110,259

(\*) Consolidated statements of financial position have been translated solely for the convenience of the reader at an exchange rate of 1USD = 115.17 JPY, the Noon Buying Rate certified by the Federal Reserve Bank of New York on December 30, 2021. The rate and methodologies used for the convenience translations differ from the currency exchange rates and translation methodologies under IFRS used for the preparation of the interim consolidated financial statements. The translation should not be construed as a representation that the Japanese yen amounts could be converted into U.S. dollars at the above or any other rate.

#### (4) Condensed Interim Consolidated Statements of Changes in Equity

Nine-month period ended December 31, 2020 (From April 1 to December 31, 2020)

	JPY (millions)					
	Equity attributable to owners of the company				Other components of equity	
	Share capital	Share premium	Treasury shares	Retained earnings	Exchange differences on translation of foreign operations	Changes in fair value of financial assets measured at fair value through other comprehensive income
As of April 1, 2020	1,668,123	1,680,287	(87,463)	1,369,972	91,848	22,891
Net profit for the period				178,907		
Other comprehensive income (loss)					(42,191)	69,348
Comprehensive income (loss) for the period	—	—	—	178,907	(42,191)	69,348
Transaction with owners:						
Issuance of new shares	22	22				
Acquisition of treasury shares			(2,138)			
Disposal of treasury shares		(0)	2			
Dividends				(283,718)		
Transfers from other components of equity				41,407		(46,286)
Share-based compensation		28,119				
Exercise of share-based awards		(29,772)	30,032			
Total transactions with owners	22	(1,631)	27,896	(242,311)	—	(46,286)
As of December 31, 2020	1,668,145	1,678,656	(59,567)	1,306,568	49,657	45,953

	Equity attributable to owners of the company				Other components of equity		
	Cash flow hedges	Hedging cost	Remeasurements of defined benefit pension plans	Total	Total	Non-controlling interests	Total equity
	As of April 1, 2020	(22,730)	555	—	92,564	4,723,483	4,003
Net profit for the period				—	178,907	120	179,027
Other comprehensive income (loss)	(21,596)	(10,288)	(4,879)	(9,606)	(9,606)	29	(9,577)
Comprehensive income (loss) for the period	(21,596)	(10,288)	(4,879)	(9,606)	169,301	149	169,450
Transaction with owners:							
Issuance of new shares				—	44		44
Acquisition of treasury shares				—	(2,138)		(2,138)
Disposal of treasury shares				—	2		2
Dividends				—	(283,718)	(77)	(283,795)
Transfers from other components of equity			4,879	(41,407)	—		—
Share-based compensation				—	28,119		28,119
Exercise of share-based awards				—	260		260
Total transactions with owners	—	—	4,879	(41,407)	(257,431)	(77)	(257,508)
As of December 31, 2020	(44,326)	(9,733)	—	41,551	4,635,353	4,075	4,639,428

Nine-month period ended December 31, 2021 (From April 1 to December 31, 2021)

JPY (millions)						
Equity attributable to owners of the company						
	Share capital	Share premium	Treasury shares	Retained earnings	Other components of equity	
					Exchange differences on translation of foreign operations	Changes in fair value of financial assets measured at fair value through other comprehensive income
As of April 1, 2021	1,668,145	1,688,424	(59,552)	1,509,906	400,798	41,983
Net profit for the period				241,417		
Other comprehensive income (loss)					206,337	(5,883)
Comprehensive income (loss) for the period	—	—	—	241,417	206,337	(5,883)
Transaction with owners:						
Issuance of new shares	8,118	14,036				
Acquisition of treasury shares			(54,451)			
Disposal of treasury shares		(0)	1			
Dividends				(284,246)		
Changes in ownership				(2,143)		
Transfers from other components of equity				1,992		(4,904)
Share-based compensation		32,057				
Exercise of share-based awards		(36,955)	22,989			
Total transactions with owners	8,118	9,138	(31,461)	(284,397)	—	(4,904)
As of December 31, 2021	1,676,263	1,697,562	(91,013)	1,466,926	607,135	31,196

Equity attributable to owners of the company							
Other components of equity							
	Cash flow hedges	Hedging cost	Remeasurements of defined benefit pension plans	Total	Total	Non-controlling interests	
						Total	Total equity
As of April 1, 2021	(68,075)	(8,592)	—	366,114	5,173,037	4,140	5,177,177
Net profit for the period				—	241,417	124	241,541
Other comprehensive income (loss)	13,958	5,969	(2,912)	217,470	217,470	33	217,503
Comprehensive income (loss) for the period	13,958	5,969	(2,912)	217,470	458,887	157	459,044
Transaction with owners:							
Issuance of new shares				—	22,154		22,154
Acquisition of treasury shares				—	(54,451)		(54,451)
Disposal of treasury shares				—	1		1
Dividends				—	(284,246)		(284,246)
Changes in ownership				—	(2,143)	(3,804)	(5,948)
Transfers from other components of equity			2,912	(1,992)	—		—
Share-based compensation				—	32,057		32,057
Exercise of share-based awards				—	(13,966)		(13,966)
Total transactions with owners	—	—	2,912	(1,992)	(300,594)	(3,804)	(304,399)
As of December 31, 2021	(54,116)	(2,623)	—	581,592	5,331,330	493	5,331,822

## (5) Consolidated Statements of Cash Flows

	JPY (millions)		USD (millions)(*)
	Nine-month Period Ended December 31,		Nine-month Period Ended December 31,
	2020	2021	2021
Cash flows from operating activities:			
Net profit for the period	¥ 179,027	¥ 241,541	\$ 2,097
Depreciation and amortization	420,281	430,877	3,741
Impairment losses	10,118	14,666	127
Equity-settled share-based compensation	28,119	32,057	278
Change in estimate of liabilities related to SHP647	(60,179)	—	—
Loss (gain) on sales and disposal of property, plant and equipment	(3,435)	258	2
Gain on divestment of business and subsidiaries	(38,273)	(1,095)	(10)
Change in fair value of financial assets and liabilities associated with contingent consideration arrangements, net	8,888	(9,683)	(84)
Finance (income) and expenses, net	115,359	100,589	873
Share of loss of investments accounted for using the equity method	8,013	5,255	46
Income tax expenses	56,330	115,077	999
Changes in assets and liabilities:			
Decrease (increase) in trade and other receivables	(49,908)	82,243	714
Decrease (increase) in inventories	6,059	(39,268)	(341)
Decrease in trade and other payables	(5,082)	(1,797)	(16)
Increase (decrease) in provisions	66,844	(70,098)	(609)
Increase (decrease) in other financial liabilities	25,939	(51,158)	(444)
Other, net	(11,810)	(858)	(7)
Cash generated from operations	756,290	848,607	7,368
Income taxes paid	(174,694)	(107,224)	(931)
Tax refunds and interest on tax refunds received	28,375	6,138	53
Net cash from operating activities	609,971	747,521	6,491
Cash flows from investing activities:			
Interest received	752	2,468	21
Dividends received	215	2,598	23
Acquisition of property, plant and equipment	(75,041)	(87,673)	(761)
Proceeds from sales of property, plant and equipment	42,818	412	4
Acquisition of intangible assets	(49,469)	(46,541)	(404)
Acquisition of investments	(9,479)	(7,600)	(66)
Proceeds from sales and redemption of investments	73,717	16,065	139
Acquisition of businesses, net of cash and cash equivalents acquired	—	(49,672)	(431)
Proceeds from sales of business, net of cash and cash equivalents divested	124,969	2,138	19
Other, net	(8,283)	(4,683)	(41)
Net cash from (used in) investing activities	100,199	(172,487)	(1,498)

	JPY (millions)		USD
	Nine-month Period Ended		Nine-month
	December 31,		Period Ended
	2020	2021	December 31,
			2021
Cash flows from financing activities:			
Net decrease in short-term loans and commercial papers	(84,997)	(2)	(0)
Proceeds from issuance of bonds and long-term loans	1,179,515	249,334	2,165
Repayments of bonds and long-term loans	(1,389,102)	(635,047)	(5,514)
Payments for settlement of forward rate agreement related to bonds	(34,830)	—	—
Acquisition of treasury shares	(2,138)	(52,538)	(456)
Interest paid	(84,185)	(84,917)	(737)
Dividends paid	(274,679)	(273,024)	(2,371)
Repayments of lease liabilities	(27,710)	(29,904)	(260)
Other, net	(156)	(366)	(3)
Net cash used in financing activities	(718,282)	(826,465)	(7,176)
Net decrease in cash and cash equivalents	(8,112)	(251,430)	(2,183)
Cash and cash equivalents at the beginning of the year			
(Consolidated statements of financial position)	637,614	966,222	8,390
Effects of exchange rate changes on cash and cash equivalents	(11,797)	9,549	83
Cash and cash equivalents at the end of the period	617,705	724,341	6,289
Cash and cash equivalents reclassified to assets held for sale	(70)	—	—
Cash and cash equivalents at the end of the period			
(Consolidated statements of financial position)	617,635	724,341	6,289

(\*) Consolidated statements of cash flows have been translated solely for the convenience of the reader at an exchange rate of 1USD = 115.17 JPY, the Noon Buying Rate certified by the Federal Reserve Bank of New York on December 30, 2021. The rate and methodologies used for the convenience translations differ from the currency exchange rates and translation methodologies under IFRS used for the preparation of the interim consolidated financial statements. The translation should not be construed as a representation that the Japanese yen amounts could be converted into U.S. dollars at the above or any other rate.



## **(6) Other Information**

### (Significant Subsequent Events)

On January 28, 2022, Takeda provided a notice of redemption to the holders of 1,500 million USD in unsecured U.S. dollar-denominated senior notes issued in September 2016 in advance of their original maturity date of September 23, 2023. The redemption date of the unsecured senior notes will be March 24, 2022. The impact from the accelerated debt prepayment on the consolidated statements of profit or loss is not expected to be material.

## Supplementary Information

<a href="#">1. Pipeline</a>	<a href="#">42</a>
• <a href="#">I. Clinical Development Activities</a>	<a href="#">42</a>
• <a href="#">II. Recent Progress in stage</a>	<a href="#">48</a>
• <a href="#">III. Discontinued projects</a>	<a href="#">49</a>
• <a href="#">IV. Main Research &amp; Development collaborations</a>	<a href="#">50</a>
<a href="#">2. Supplementary Financial Information</a>	<a href="#">56</a>
• <a href="#">Revenue by region</a>	<a href="#">56</a>
◦ <a href="#">Year to date</a>	<a href="#">56</a>
◦ <a href="#">Quarterly</a>	<a href="#">57</a>
• <a href="#">Product Sales Analysis</a>	<a href="#">58</a>
◦ <a href="#">Year to date</a>	<a href="#">58</a>
◦ <a href="#">Quarterly</a>	<a href="#">60</a>
■ <a href="#">Q1</a>	<a href="#">60</a>
■ <a href="#">Q2</a>	<a href="#">62</a>
■ <a href="#">Q3</a>	<a href="#">64</a>
• <a href="#">Product Sales Analysis (Reported &amp; Underlying Growth)</a>	<a href="#">66</a>
• <a href="#">Product Forecast</a>	<a href="#">68</a>
• <a href="#">Exchange Rate</a>	<a href="#">70</a>
• <a href="#">CAPEX, depreciation and amortization and impairment losses</a>	<a href="#">71</a>
<a href="#">3. Reconciliation</a>	<a href="#">72</a>
• <a href="#">FY2021 Q3 YTD Reconciliation from Reported Revenue to Core/Underlying Revenue</a>	<a href="#">72</a>
• <a href="#">FY2021 Q3 YTD Reconciliation from Reported to Core/Underlying Core</a>	<a href="#">73</a>
• <a href="#">FY2020 Q3 YTD Reconciliation from Reported to Core/Underlying Core</a>	<a href="#">74</a>
• <a href="#">Free Cash Flow</a>	<a href="#">75</a>
• <a href="#">FY2021 Q3 YTD LTM Net Profit to Adjusted EBITDA Bridge</a>	<a href="#">76</a>
• <a href="#">FY2021 Q3 YTD Net Debt to Adjusted EBITDA</a>	<a href="#">77</a>
• <a href="#">FY2020 Net Debt to Adjusted EBITDA</a>	<a href="#">78</a>
• <a href="#">Reconciliation from Reported Operating Profit to Core Operating Profit - FY2021 Forecast</a>	<a href="#">79</a>

# 1. Pipeline

## I. Clinical Development Activities

- The following table lists the pipeline assets that we are developing as of February 3, 2022. The assets in our pipeline are in various stages of development, and the contents of the pipeline may change as compounds currently under development drop out and new compounds are introduced. Whether the compounds listed below are ever successfully released as products depends on various factors, including the results of pre-clinical and clinical trials, market conditions for various drugs and regulatory approvals.
- This table primarily shows the indications for which we will actively pursue approval. We are also conducting additional studies of certain assets to examine their potential for use in further indications and in additional formulations.
- The listings in this table are limited to the U.S., EU and Japan and China, but we are also actively conducting development activities in other regions, including in Emerging Markets. Country/region column denote where a pivotal clinical study is ongoing or a filing has been made with our specific intention to pursue approval in any of the U.S., EU, Japan or China. 'Global' refers to U.S., EU, Japan and China.
- Brand name and country/region indicate the brand name and country in which the specific asset has already been approved for any indication in any of the U.S., EU, Japan or China and Takeda has commercialization rights for such asset.
- Stage-ups are recognized in the table upon achievement of First Subject In.
- Modality of our pipeline assets in the following table is classified into either of the following categories: 'small molecule', 'peptide/oligonucleotide', 'cell and gene therapy', 'microbiome' or 'biologic and other.'

### • Oncology Pipeline

Development code <generic name> Brand name (country/region)	Type of Drug (administration route)	Modality	Indications / additional formulations	Country /Region	Stage
<b>SGN-35<sup>1</sup></b> <brentuximab vedotin> <i>ADCESTRIS</i> (EU, Japan, China)	CD30 monoclonal antibody-drug conjugate (injection)	Biologic and other	Cutaneous T cell lymphoma	China	Approved (Apr 2021)
<brigatinib> <i>ALUNBRIG</i> (Global)	ALK inhibitor (oral)	Small molecule	1L & 2L ALK-positive Non-Small Cell Lung Cancer	China	Filed (Dec 2020)
			2L ALK-positive Non-Small Cell Lung Cancer (head-to-head with alectinib)	Global	P-III
<b>MLN9708</b> <ixazomib> <i>NINLARO</i> (Global)	Proteasome inhibitor (oral)	Small molecule	Maintenance therapy in patients with newly diagnosed Multiple Myeloma not treated with stem cell transplant	Japan U.S. EU China	Approved (May 2021) P-III P-III P-III
			Maintenance therapy in patients with newly diagnosed Multiple Myeloma following autologous stem cell transplant	U.S. EU	P-III P-III
<cabozantinib> <sup>2</sup> <i>CABOMETYX</i> (Japan)	Multi-targeted kinase inhibitor (oral)	Small molecule	1L Renal cell carcinoma in combination with nivolumab	Japan	Approved (Aug 2021)
			2L metastatic Non-Small Cell Lung Cancer in combination with atezolizumab <sup>3</sup>	Japan	P-III
			Metastatic Castration-Resistant Prostate Cancer in combination with atezolizumab <sup>4</sup>	Japan	P-III
<ponatinib> <i>ICLUSIG</i> (U.S.)	BCR-ABL inhibitor (oral)	Small molecule	Front line Philadelphia chromosome-positive Acute Lymphoblastic Leukemia	U.S.	P-III
<b>TAK-788</b> <mobocertinib> <i>EXKIVITY</i> (U.S.)	EGFR/HER2 exon 20 inhibitor (oral)	Small molecule	Treatment Naïve Non-Small Cell Lung Cancer with EGFR exon 20 insertion	Global	P-III
			Previously treated Non-Small Cell Lung Cancer with EGFR exon 20 insertion <sup>5</sup>	U.S.	Approved (Sep 2021)
				China	Filed (Jul 2021)
				Japan EU	P-III P-III
<b>TAK-385</b> <relugolix>	LH-RH antagonist (oral)	Small molecule	Prostate cancer	Japan	P-III
				China	P-III

[Table of Contents](#)

<b>TAK-981</b> <subasumstat>	SUMO inhibitor (injection)	Small molecule	Multiple cancers	-	P-II
<b>TAK-007</b> <sup>6</sup>	CD19 CAR-NK (injection)	Cell and gene therapy	Relapsed/refractory B cell malignancies	-	P-I/II
<b>TAK-102</b> <sup>7</sup>	GPC3 CAR-T (injection)	Cell and gene therapy	Solid tumors	-	P-I
<b>TAK-573</b> <sup>8</sup> <modakafusp alfa>	Anti-CD38-targeted IgG4 genetically fused with an attenuated IFN $\alpha$ (injection)	Biologic and other	Relapsed/refractory Multiple Myeloma	-	P-I
<b>TAK-605</b> <sup>9</sup>	Oncolytic virus (intra-tumoral administration)	Biologic and other	Solid tumors	-	P-I
<b>TAK-676</b>	STING agonist (injection)	Small molecule	Solid tumors	-	P-I
<b>TAK-940</b> <sup>10</sup>	CD19 1XX CAR-T (injection)	Cell and gene therapy	Relapsed/refractory B cell malignancies	-	P-I
<b>TAK-186</b> <sup>11</sup>	T Cell Engager (Injection)	Biologic and other	EGFR expressing solid tumors	-	P-I

1. Partnership with Seagen, Inc.
2. Partnership with Exelixis, Inc.
3. Partnership with Chugai Pharmaceutical. Chugai operates Phase 3 development
4. Partnership with Chugai Pharmaceutical. Takeda operates Phase 3 development
5. The U.S. FDA review is being conducted under Project Orbis, an initiative of the FDA Oncology Center of Excellence (OCE), which provides a framework for concurrent submission and review of oncology products among international partners such as the UK, Brazil and Australia.
6. Partnership with The University of Texas MD Anderson Cancer Center
7. Partnership with Noile-Immune Biotech, Inc.
8. Partnership with Teva Pharmaceutical Industries Ltd.
9. Partnership with Turnstone Biologics
10. Partnership with Memorial Sloan Kettering Cancer Center
11. Acquired via acquisition of Maverick Therapeutics, Inc.

Additions since FY2021 Q2: None

Removals since FY2021 Q2: TAK-252 for solid tumors or lymphomas (P-I, collaboration agreement with Shattuck Labs terminated)

• Rare Genetics and Hematology Pipeline

Development code <generic name> Brand name (country/region)	Type of Drug (administration route)	Modality	Indications / additional formulations	Country /Region	Stage
<b>TAK-743</b> <lanadelumab> <i>TAKHZYRO</i> (U.S., EU, China)	Plasma kallikrein inhibitor (injection)	Biologic and other	Hereditary Angioedema	Japan	Filed (Mar 2021)
			Pediatric Hereditary Angioedema	Global	P-III
			Bradykinin-Mediated Angioedema	Global	P-III
<b>TAK-577</b> <i>VONVENDI</i> (U.S., Japan), <i>VEYVONDI</i> (EU)	von Willebrand factor [recombinant] (injection)	Biologic and other	Adult prophylactic treatment of von Willebrand disease	U.S. Japan EU China	Approved (Jan 2022)* P-III P-III P-III
			Pediatric on-demand and surgery treatment of von Willebrand disease	Global	P-III
<b>TAK-660</b> <i>ADYNOVATE</i> (U.S., Japan) <i>ADYNOVI</i> (EU)	Antihemophilic factor [recombinant], PEGylated (injection)	Biologic and other	Pediatric Hemophilia A	EU	P-III
<b>TAK-755</b> <sup>1</sup>	Replacement of the deficient-ADAMTS13 enzyme (injection)	Biologic and other	Congenital Thrombotic Thrombocytopenic Purpura	U.S. EU	P-III P-III
			Immune Thrombotic Thrombocytopenic Purpura	U.S. EU	P-II P-II
			Sickle cell disease	U.S.	P-I
<b>TAK-620</b> <sup>2</sup> <maribavir> <i>LIVTENCITY</i> (U.S.)	Benzimidazole riboside inhibitor (oral)	Small molecule	Post-transplant CMV infection/disease resistant/refractory to (val) ganciclovir, cidofovir or foscarnet	U.S. EU	Approved (Nov 2021) Filed (Jun 2021)
			HSCT Recipients with First CMV Infection	U.S. EU	P-III P-III
<b>TAK-609</b>	Recombinant human iduronate-2-sulfatase for intrathecal administration (injection)	Biologic and other	Hunter syndrome CNS	U.S. EU	P-II P-II
<b>TAK-611</b>	Recombinant human arylsulfatase A for intrathecal administration (injection)	Biologic and other	Metachromatic leukodystrophy	-	P-II
<b>TAK-079</b> <sup>3</sup> <mezagitamab>	Anti-CD38 monoclonal antibody (injection)	Biologic and other	Myasthenia gravis	-	P-II
			Immune thrombocytopenic purpura	-	P-II
			Systemic lupus erythematosus	-	P-I/II
<b>TAK-834</b> <i>NATPARA</i> (U.S.), <i>NATPAR</i> (EU)	Parathyroid hormone (injection)	Biologic and other	Hypoparathyroidism	Japan	P-I <sup>4</sup>

1. Partnership with KM Biologics for co-exclusive license for commercialization in Japan only
2. Partnership with GlaxoSmithKline
3. Relapsed/refractory Multiple Myeloma will continue until trial completion.
4. P-I study in Japan completed; P-III study start timing under review.

\* Event after the Q3 reporting period: Update after January 1, 2022

Additions since FY2021 Q2: None

Removals since FY2021 Q2: Tak-607 for Complications of prematurity (P-II, externalization)

• **Neuroscience Pipeline**

Development code <generic name> Brand name (country/region)	Type of Drug (administration route)	Modality	Indications / additional formulations	Country /Region	Stage
<b>TAK-935</b> <socticlestat>	CH24H inhibitor (oral)	Small molecule	Dravet syndrome	-	P-III
			Lennox-Gastaut syndrome	-	P-III
<b>TAK-994</b>	Orexin 2R agonist (oral)	Small molecule	Narcolepsy	-	P-II <sup>3</sup>
<b>TAK-071</b>	M1 positive allosteric modulator (M1PAM) (oral)	Small molecule	Parkinson's disease	-	P-II
<b>TAK-041</b> <sup>1</sup>	GPR139 agonist (oral)	Small molecule	Anhedonia in major depressive disorder (MDD)	-	P-II
<b>TAK-341/MEDI1341</b> <sup>2</sup>	Alpha-synuclein antibody (injection)	Biologic and other	Parkinson's disease	-	P-I
<b>TAK-653</b> <sup>1</sup>	AMPA receptor potentiator (oral)	Small molecule	Inadequate response to treatment in major depressive disorder (MDD)	-	P-I
<b>TAK-861</b>	Orexin 2R agonist (oral)	Small molecule	Sleep disorders, other disorders	-	P-I
<b>TAK-925</b>	Orexin 2R agonist (injection)	Small molecule	Hospital setting, narcolepsy	-	P-I

1. 50:50 co-development and co-commercialization with Neurocrine
2. Partnership with AstraZeneca. AstraZeneca leads Phase 1 development
3. TAK-994 currently on clinical hold in the U.S.

Additions since FY2021 Q2: None  
Removals since FY2021 Q2: None

• **GI Pipeline**

Development code <generic name> Brand name (country/region)	Type of Drug (administration route)	Modality	Indications / additional formulations	Country /Region	Stage
<b>MLN0002</b> <vedolizumab> <i>ENTYVIO</i> (Global)	Humanized monoclonal antibody against $\alpha 4\beta 7$ integrin (injection)	Biologic and other	Subcutaneous formulation for ulcerative colitis	U.S.	Complete Response Letter (CRL) received (Dec 2019) <sup>7</sup>
				Japan	Filed (Aug 2019)
			Subcutaneous formulation for Crohn's disease	U.S.	P-III
				Japan	P-III
			Active Chronic Pouchitis	EU	Approved (Jan 2022)*
			Graft-versus-Host Disease prophylaxis in patients undergoing allogeneic hematopoietic stem cell transplantation	EU Japan	P-III P-III
		Pediatrics Study (ulcerative colitis, Crohn's disease)	Global	P-II	
<b>TAK-438</b> <vonoprazan> <i>TAKECAB</i> (Japan) <i>VOCINTI</i> (China)	Potassium-competitive acid blocker (oral)	Small molecule	Acid related diseases (Reflux Esophagitis Maintenance)	China	Approved (Oct 2021)
			Oral disintegrated tablet formulation	Japan	Filed (Mar 2021)
			Acid related diseases (adjunct to <i>Helicobacter pylori</i> eradication)	China	P-III

[Table of Contents](#)

<b>TAK-633</b> <b>&lt;teduglutide&gt;</b> <i>GATTEX</i> (U.S.) <i>REVESTIVE</i> (EU, Japan)	GLP-2 analogue (injection)	Peptide/ Oligo- nucleotide	Short bowel syndrome (pediatric indication)	Japan	Approved (Jun 2021)
			Short bowel syndrome (in adults)	Japan	Approved (Jun 2021)
<b>Cx601</b> <b>&lt;darvadstrocel&gt;</b> <i>ALOFISEL</i> (EU, Japan)	A suspension of allogeneic expanded adipose- derived stem cells (injection)	Biologic and other	Refractory complex perianal fistulas in patients with Crohn's disease	U.S. Japan	P-III Approved (Sep 2021)
<b>TAK-906</b>	Dopamine D2/D3 receptor antagonist (oral)	Small molecule	Gastroparesis	-	P-II (b)
<b>TAK-954</b> <sup>1</sup>	5-HT <sub>4</sub> - hydroxytryptamine receptor agonist (injection)	Small molecule	Post-operative gastrointestinal dysfunction	-	P-II (b)
<b>TAK-999</b> <sup>2</sup>	GalNAc based RNA interference (RNAi) (injection)	Peptide/ Oligo- nucleotide	Alpha-1 antitrypsin-deficiency associated liver disease	U.S. EU	P-II (b) P-II (b)
<b>TAK-101</b> <sup>3</sup>	Tolerizing Immune Modifying nanoParticle (TIMP) (injection)	Biologic and other	Celiac disease	-	P-II (a)
<b>TAK-018/EB8018</b> <sup>4</sup> <b>&lt;sibofimloc&gt;</b>	FimH antagonist (oral)	Small molecule	Crohn's disease (post-operative and ileal-dominant)	-	P-II (a)
<b>TAK-951</b>	Peptide agonist (sub- cutaneous)	Peptide/ Oligo- nucleotide	Nausea and vomiting	-	P-II
<b>TAK-510</b>	Peptide agonist (sub- cutaneous)	Peptide/ Oligo- nucleotide	Nausea and vomiting	-	P-I
<b>TAK-105</b>	Peptide agonist (sub- cutaneous)	Peptide/ Oligo- nucleotide	Nausea and vomiting	-	P-I
<b>TAK-062</b>	Glutenase (oral)	Biologic and other	Celiac disease	-	P-I
<b>TAK-039</b> <sup>5</sup>	Bacterial consortium (oral)	Microbiome	Clostridium difficile infections <sup>6</sup>	-	P-I

1. Partnership with Theravance Biopharma, Inc.

2. Partnership with Arrowhead Pharmaceuticals, Inc.

3. Acquired development and commercialization license for TAK-101 from Cour Pharmaceutical Development Company. Previously known as TIMP-GLIA.

4. Partnership with Enterome Bioscience SA

5. Partnership with NuBiyota

6. Phase 1 study in clostridium difficile infections completed; strategic intention is to take the program forward in hepatic encephalopathy.

7 In active discussions with the FDA. Timelines under review; potential approval anticipated FY2023.

\* Event after the Q3 reporting period: Update after January 1, 2022

Additions since FY2021 Q2: None

Removals since FY2021 Q2: TAK-438 for Acid related diseases (Duodenal Ulcer) (China - Filing withdrawn, discontinued)

TAK-721 for Eosinophilic esophagitis (U.S. - Filing, discontinued)

• **Plasma-Derived Therapies Pipeline**

Development code <generic name> Brand name (country/region)	Type of Drug (administration route)	Modality	Indications / additional formulations	Country/ Region	Stage
<b>TAK-664</b> <i>CUVITRU</i> (U.S., EU)	Immunoglobulin 20% [human] (subcutaneous)	Biologic and other	Primary immunodeficiencies	Japan	P-III
<b>TAK-771<sup>1</sup></b> < <b>IG Infusion 10%</b> ( <b>Human</b> ) w/ <b>Recombinant Human</b> <b>Hyaluronidase</b> > <i>HYQVIA</i> (U.S., EU)	Immunoglobulin (IgG) + recombinant hyaluronidase replacement therapy (injection)	Biologic and other	Pediatric indication for primary immunodeficiency	U.S.	P-III
			Chronic inflammatory demyelinating polyradiculoneuropathy	U.S. EU	P-III P-III
			Chronic inflammatory demyelinating polyradiculoneuropathy and Multifocal Motor Neuropathy	Japan	P-III*
<b>TAK-662</b> <i>CEPROTIN</i> (U.S., EU)	Protein C concentrate [human] (injection)	Biologic and other	Severe congenital protein C deficiency	Japan	P-I/II
<b>TAK-881</b> < <b>Facilitated 20%</b> <b>SCIG</b> >	Immunoglobulin (20%) [human] + recombinant hyaluronidase replacement therapy (injection)	Biologic and other	Immunodeficiencies	-	P-I

1. Partnership with Halozyme

\* Event after the Q3 reporting period: Update after January 1, 2022

Additions since FY2021 Q2: TAK-771 for Chronic inflammatory demyelinating polyradiculoneuropathy and Multifocal Motor Neuropathy (Japan, P-III)  
TAK-881 for Immunodeficiencies (P-I)

Removals since FY2021 Q2: None

• **Vaccines Pipeline**

Development code Brand name (country/region)	Type of vaccine (administration route)	Modality	Indications / additional formulations	Country/ Region	Stage
<b>TAK-919/mRNA-1273<sup>1</sup></b> <i>Spikevax Intermuscular Injection</i> (Japan)	SARS-CoV-2 vaccine (injection)	Biologic and other	Active immunization for the prevention of COVID-19	Japan	Approved (May 2021) <sup>4</sup>
			Active immunization for the prevention of COVID-19 (booster)	Japan	Approved (Dec 2021)
<b>TAK-003</b>	Tetravalent dengue vaccine (injection)	Biologic and other	For the prevention of dengue fever of any severity, due to any serotype, in individuals aged 4 up to 60 years of age	EU and EU- M4all -	Filed (Mar 2021) <sup>5</sup>  P-III
<b>TAK-019/ NVX-CoV2373<sup>2</sup></b>	SARS-CoV-2 vaccine (injection)	Biologic and other	Active immunization for the prevention of COVID-19	Japan	Filed (Dec 2021)
<b>TAK-426<sup>3</sup></b>	Zika vaccine (injection)	Biologic and other	Active immunization for the prevention of disease caused by Zika virus	-	P-I

1. Partnership with Moderna and MHLW

2. Partnership with Novavax, Inc.

3. Partnership with The Biomedical Advanced Research and Development Authority (BARDA) of the U.S. Government

4. Change in age indication to expand to 12 years of age and older (July 2021).

5. In addition to filing in the EU and through the EU-M4all (previously Article 58) procedure for countries outside of the EU, filings began in dengue endemic countries in Latin America and Asia that are not participating in the EU-M4all procedure.

Additions since FY2021 Q2: TAK-919 for active immunization for the prevention of COVID-19 (booster) (Japan, Approved)

Removals since FY2021 Q2: None



**II. Recent Progress in stage [Progress in stage since April 1st, 2021]**

Development code <generic name>	Indications / additional formulations	Country/Region	Progress in stage
SGN-35 <brentuximab vedotin>	Cutaneous T cell lymphoma	China	Approved (Apr 2021)
MLN9708 <ixazomib>	Maintenance therapy in patients with newly diagnosed Multiple Myeloma not treated with stem cell transplant	Japan	Approved (May 2021)
TAK-919/mRNA-1273	Active immunization for the prevention of COVID-19	Japan	Approved (May 2021)
TAK-633 <teduglutide>	Short bowel syndrome (pediatric indication and in adults)	Japan	Approved (Jun 2021)
<cabozantinib>	IL Renal cell carcinoma in combination with nivolumab	Japan	Approved (Aug 2021)
TAK-788 <mobocertinib>	Previously treated Non-Small Cell Lung Cancer with EGFR exon 20 insertion	U.S.	Approved (Sep 2021)
Cx601 <darvadstrocel>	Refractory complex perianal fistulas in patients with Crohn's disease	Japan	Approved (Sep 2021)
TAK-438 <vonoprazan>	Acid related diseases (Reflux Esophagitis Maintenance)	China	Approved (Oct 2021)
TAK-620 <maribavir>	Post-transplant CMV infection/disease resistant/refractory to (val) ganciclovir, cidofovir or foscarnet	U.S.	Approved (Nov 2021)
TAK-919/mRNA-1273	Active immunization for the prevention of COVID-19 (booster)	Japan	Approved (Dec 2021)
TAK-577	Adult prophylactic treatment of von Willebrand disease	U.S.	Approved (Jan 2022)*
MLN0002 <vedolizumab> <vedolizumab>	Active Chronic Pouchitis	EU	Approved (Jan 2022)*
TAK-620 <maribavir>	Post-transplant CMV infection/disease resistant/refractory to (val) ganciclovir, cidofovir or foscarnet	EU	Filed (Jun 2021)
TAK-788 <mobocertinib>	Previously treated Non-Small Cell Lung Cancer with EGFR exon 20 insertion	China	Filed (Jul 2021)
TAK-019/NVX-CoV2373	Active immunization for the prevention of COVID-19	Japan	Filed (Dec 2021)
TAK-935 <soticlestat>	Dravet Syndrome	-	P-III
TAK-935 <soticlestat>	Lennox-Gastaut syndrome	-	P-III
TAK-771 <IG Infusion 10% (Human) w/ Recombinant Human Hyaluronidase>	Chronic inflammatory demyelinating polyradiculoneuropathy and Multifocal Motor Neuropathy	Japan	P-III*
TAK-981	Multiple cancers	-	P-II
TAK-041	Anhedonia in major depressive disorder (MDD)	-	P-II
TAK-662	Severe congenital protein C deficiency	Japan	P-I/II
TAK-861	Sleep disorders, other disorders	-	P-I
TAK-105	Nausea and vomiting	-	P-I
TAK-881 <Facilitated 20% SCIG>	Immunodeficiencies	-	P-I

\* Event after the Q3 reporting period: Update after January 1, 2022

**III. Discontinued projects [Update since April 1st, 2021]**

Development code <generic name>	Indications (Region/Country, Stage)	Reason
<b>CoVIg-19</b>	Treatment of adult hospitalized patients at onset of clinical progression of COVID-19 (U.S., EU, Japan, P-III)	Phase 3 Inpatient Treatment with Anti-Coronavirus Immunoglobulin (ITAC) clinical trial sponsored and funded by the National Institute of Allergy and Infectious Diseases (NIAID), part of the National Institutes of Health (NIH), did not meet its endpoints.
<b>TAK-169</b>	Relapse/refractory multiple myeloma (P-I)	Takeda has communicated its decision to turn over full rights of TAK-169 to Molecular Templates. Molecular Templates will continue to develop TAK-169.
<b>TAK-831</b> <luvadaxistat>	Negative symptoms and/or cognitive impairment associated with schizophrenia (P-II)	Based on clinical data, our partner Neurocrine announced the discontinuation of development in Schizophrenia Negative Symptoms. Neurocrine will continue developing TAK-831 in Cognitive Impairment Associated with Schizophrenia and Takeda decided not to co-fund a supplemental study with Neurocrine, which resulted in Takeda's maintaining its right to receive milestones and royalties regarding TAK-831.
<b>TAK-671</b>	Acute Pancreatitis (P-I)	Takeda has opted out of further development based on a business decision, and the right to continue developing the asset falls under Samsung Bioepis.
<b>TAK-924</b> <pevonedistat>	High-risk Myelodysplastic Syndrome (P-III), Unfit Acute Myelogenous Leukemia (P-III)	Phase 3 PANTHER study did not meet its primary endpoint. The result did not support further development in Phase 3 HR MDS trial and Unfit AML trial. The Phase 1/2 AML trial in combination with venetoclax is ongoing but not recruiting new patients and is not registrational.
<b>TAK-935</b> <sothicestat>	15q duplication syndrome, CDKL5 deficiency disorder (P-II)	The Phase 2 result did not support further development in these indications.
<b>TAK-252</b>	Solid tumors and lymphomas (P-I)	Shattuck Labs and Takeda mutually agreed to terminate the parties' Collaboration Agreement, resulting in termination of the TAK-252 for Takeda.
<b>TAK-438</b> <vonoprazan>	Acid related diseases (Duodenal Ulcer) (China, Filing withdrawn)	After evaluation of Chinese CDE (Center for Drug Evaluation) assessment, Takeda decided not to pursue this indication further.
<b>TAK-721</b> <budesonide>	Eosinophilic esophagitis (U.S., Filed)	After evaluation of Complete Response Letter (CRL) from U.S. FDA, Takeda decided not to pursue program further.

**IV. Main Research & Development collaborations [not a comprehensive list of all Takeda R&D collaborations]**

• **Oncology**

Partner	Country of incorporation	Subject
Adimab	U.S.	Agreement for the discovery, development and commercialization of three mAbs and three CD3 Bi-Specific antibodies for oncology indications.
ASKA Pharmaceutical Co., Ltd	Japan	Takeda granted exclusive commercialization rights for uterine fibroids and exclusive development and commercialization rights for endometriosis for Japan to maximize the product value of relugolix (TAK-385).
Crescendo Biologics	U.K.	Collaboration and licensing agreement for the discovery, development and commercialization of Humabody®-based therapeutics for cancer indications.
Egle Therapeutics	France	Identify novel tumor-specific regulatory T cell targets and develop unique anti-suppressor-based immunotherapies.
Exelixis, Inc.	U.S.	Exclusive licensing agreement to commercialize and develop novel cancer therapy cabozantinib and all potential future cabozantinib indications in Japan, including advanced renal cell carcinoma and hepatocellular carcinoma.
GammaDelta Therapeutics <sup>‡</sup>	U.K.	Collaboration agreement to discover and develop new immunotherapies in oncology using GammaDelta Therapeutics' novel T cell platform based on the unique properties of gamma delta T cells derived from human tissues. Takeda exercised its option to acquire GammaDelta Therapeutics in October 2021. Separately, in January 2022, Takeda exercised its option to acquire Adaptate Biotherapeutics, a UK based spin-out company from GammaDelta Therapeutics focused on developing antibody-based therapeutics for the modulation of variable delta 1 (Vδ1) gamma delta (γδ).
GlaxoSmithKline	U.K.	Exclusive licensing agreement to develop and commercialize novel cancer therapy niraparib for the treatment of all tumor types in Japan, and all tumor types excluding prostate cancer in South Korea and Taiwan.
Heidelberg Pharma	Germany	Antibody-Drug-Conjugate (ADC) research collaboration on 2 targets and licensing agreement (α-amanitin payload and proprietary linker).
KSQ Therapeutics	U.S.	Strategic collaboration to research, develop and commercialize novel immune-based therapies for cancer using KSQ's CRISPRomics® technology.
MD Anderson Cancer Center	U.S.	Exclusive license and research agreement to develop cord blood-derived chimeric antigen receptor-directed natural killer (CAR NK) cell therapies, 'armored' with IL-15, for the treatment of B cell malignancies and other cancers.
Memorial Sloan Kettering Cancer Center	U.S.	Strategic research collaboration and license to develop novel chimeric antigen receptor T cell (CAR-T) products for the treatment of multiple myeloma, acute myeloid leukemia and additional solid tumor indications. The collaboration is co-led by Michel Sadelain, who is currently head of the Center for Cell Engineering at Memorial Sloan Kettering
Molecular Templates	U.S.	Research collaboration to apply Molecular Templates' engineered toxin bodies (ETB) technology platform to potential therapeutic targets provided by Takeda, who has rights to exercise exclusive options to obtain license rights to products resulting from the collaboration.
Myovant Sciences	Switzerland	Takeda granted Myovant an exclusive, worldwide license (excluding Japan and certain other Asian countries) to relugolix (TAK-385) and an exclusive, worldwide license to MVT-602 (TAK-448).
National Cancer Center of Japan	Japan	Partnership agreement to develop basic research to clinical development by promoting exchanges among researchers, physicians, and others engaged in anti-cancer drug discovery and cancer biology research.
Noile-Immune Biotech	Japan	Collaboration agreement for the development of next generation CAR-T cell therapy, developed by Professor Koji Tamada at Yamaguchi University. Takeda has exclusive options to obtain licensing rights for the development and commercialization of Noile-Immune Biotech's pipeline and products resulting from this partnership. Due to the success of the collaboration, Takeda licensed NIB-102 and NIB-103.
Presage Biosciences	U.S.	Research collaboration and license for multiple programs using Presage's proprietary platform CIVO to evaluate patients' unique responses to microdoses of cancer drugs.
Seagen	U.S.	Agreement for the joint development of ADCETRIS, an ADC technology which targets CD30 for the treatment of HL. Approved in 67 countries with ongoing clinical trials for additional indications.
Teva	Israel	Agreement for worldwide License to TEV-48573 (TAK-573) (modakafusp alfa, Anti-CD38-Attenukine™) and multi-target discovery collaboration accessing Teva's attenukine platform.
Turnstone Biologics	U.S.	Collaboration to co-develop TAK-605 (RIVAL-01) (novel oncolytic virus expressing aCTLA4, IL12-mb, flt3L) via a worldwide partnership and also conduct collaborative discovery efforts to identify additional novel product candidates based on a Turnstone's vaccinia virus platform.

<sup>‡</sup> Executed since April 1, 2021

**Rare Genetics and Hematology**

Partner	Country of incorporation	Subject
Asklepios Biopharmaceuticals	U.S.	Agreement for multiple research and development collaborations using FVIII Gene Therapy for the treatment of Hemophilia A and B.
BioMarin	U.S.	Agreement for the in-license of enabling technology for the exogenous replacement of iduronate-2-sulfatase with Idursulfase-IT in patients via direct delivery to the CNS for the long-term treatment of Hunter Syndrome in patients with cognitive impairment in order to slow progression of cognitive impairment (TAK-609).
Carmine Therapeutics	Singapore	Research collaboration agreement to discover, develop and commercialize transformative non-viral gene therapies for two rare disease targets using Carmine's REGENT(TM) technology, based on red blood cell extracellular vesicles.
Codexis, Inc.	U.S.	Strategic collaboration and license for the research and development of novel gene therapies for certain disease indications, including the treatment of lysosomal storage disorders and blood factor deficiencies.
Ensoma	U.S.	Research collaboration and license provides Takeda with an exclusive worldwide license to Ensoma's Engenius™ vectors for up to five rare disease indication.
Evox Therapeutics	U.K.	Collaboration for developing novel protein replacement and mRNA therapies and targeted delivery using Evox's proprietary exosome technology. Partnership for up to five rare disease targets with Takeda assuming responsibility for its clinical development
GlaxoSmithKline	U.K.	In-license agreement between GSK and University of Michigan for TAK-620 (marabivir) in the treatment of human cytomegalovirus.
JCR Pharmaceuticals‡	Japan	Exclusive collaboration and license agreement to commercialize JR-141 (pabinafusp alfa), applied with J-Brain Cargo®, JCR's proprietary blood-brain barrier (BBB) technology, for the treatment of Hunter syndrome (MPS II). Takeda will exclusively commercialize JR-141 outside of the United States, including Canada, Europe, and other regions (excluding Japan and certain other Asia-Pacific countries). Takeda receives an option under a separate option agreement, which allows Takeda to acquire an exclusive license to commercialize JR-141 in the U.S. upon completion of the Phase 3 program.
Immusoft‡	U.S.	Research collaboration and license option agreement to discover, develop and commercialize cell therapies in rare inherited metabolic disorders with central nervous system (CNS) manifestations and complications using Immusoft's Immune System Programming (ISP™) technology platform.
IPSEN	France	Purchase agreement for the development of Obizur for the treatment of Acquired Hemophilia A including for patients with Congenital Hemophilia A with inhibitors indication in elective or emergency surgery.
KM Biologics	Japan	Agreement for the development collaboration of TAK-755 to overcome the ADAMTS13 deficiency in TTP.
Oak Hill Bio‡*	UK	Multiple asset and license agreements with Oak Hill Bio, a rare disease therapeutics company. Takeda transfers multiple pre-clinical and clinical programs, including OHB-607 (formerly TAK-607) and OHB-101 (formerly TAK-752), to Oak Hill Bio in exchange for an upfront payment, an ownership stake in Oak Hill Bio and potential milestones and royalty payments.
Poseida Therapeutics‡	U.S.	Research collaboration and exclusive license agreement to utilize Poseida's piggyBac, Cas-CLOVER, biodegradable DNA and RNA nanoparticle delivery technology and other proprietary genetic engineering platforms for up to eight gene therapies. The collaboration will focus on developing non-viral in vivo gene therapy programs, including Poseida's Hemophilia A program.
Selecta Biosciences‡	U.S.	Research collaboration and license agreement to develop targeted, next-generation gene therapies for two indications within the field of lysosomal storage disorders using Selecta's ImmTOR platform.
Xenetic Biosciences	U.S.	Exclusive R&D license agreement for PolyXen delivery technology for hemophilia factors VII, VIII, IX, X.

‡ Executed since April 1, 2021

\* Event after the Q3 reporting period: Update after January 1, 2022

• **Neuroscience**

Partner	Country of incorporation	Subject
Anima Biotech	U.S.	Strategic collaboration to discover and develop mRNA translation modulators for genetically-defined neurological diseases.
AstraZeneca	UK	Agreement for the joint development and commercialization of MEDI1341, an alpha-synuclein antibody currently in development as a potential treatment for Parkinson’s disease.
BridGene Biosciences	U.S.	Research collaboration to discover small molecule drugs for “undruggable” targets using BridGene’s chemoproteomics platform.
CNDAP (Cure Network Dolby Acceleration Partners)*	U.S.	Research collaboration to develop small molecules targeting tau, a protein involved in Alzheimer’s disease and other major brain disorders.
Denali Therapeutics	U.S.	Strategic option and collaboration agreement to develop and commercialize up to three specified therapeutic product candidates for neurodegenerative diseases, incorporating Denali’s transport vehicle (TV) platform for increased exposure of biotherapeutic products in the brain; options exercised on DNL593/TAK-594 and DNL919/TAK-920 in Q3 FY2021.
Lundbeck	Denmark	Collaboration agreement to develop and commercialize vortioxetine.
Neurocrine Biosciences	U.S.	Collaboration to develop and commercialize 7 compounds in Takeda’s early-to-mid stage neuroscience pipeline, including TAK-041, TAK-653 and TAK-831. Takeda will be entitled to certain development milestones, commercial milestones and royalties on net sales and will, at certain development events, be able opt in or out of a 50:50 profit share on all clinical programs on an asset-by-asset basis.
PeptiDream‡	Japan	Collaborative research and exclusive license agreement to create peptide-drug conjugates (PDCs) for neuromuscular and neurodegenerative diseases.
Skyhawk Therapeutics	U.S.	Collaboration and licensing agreement to develop and commercialize RNA modulation therapies targeting neurodegenerative diseases.
StrideBio	U.S.	Collaboration and license agreement to develop <i>in vivo</i> adeno-associated viruses (AAV) based therapies for Friedreich’s Ataxia (FA) and two additional undisclosed targets.
Wave Life Sciences	Singapore	Multi-program option agreement to co-develop and co-commercialize antisense oligonucleotides for a range of neurological diseases.

‡ Executed since April 1, 2021

• **Gastroenterology**

Partner	Country of incorporation	Subject
Ambys Medicines	U.S.	Collaboration agreement for the application of novel modalities, including cell and gene therapy and gain-of-function drug therapy, to meet the urgent need for treatments that restore liver function and prevent the progression to liver failure across multiple liver diseases. Under the terms of the agreement, Takeda has an option to ex-U.S. commercialization rights for the first 4 products that reach an investigational new drug application.
Arcturus	U.S.	Collaboration agreement to develop RNA-based therapeutics for the treatment of non-alcoholic steatohepatitis and other gastrointestinal related disorders using Arcturus’ wholly-owned LUNAR™ lipid-mediated delivery systems and UNA Oligomer chemistry.
Arrowhead Pharmaceuticals	U.S.	Collaboration and licensing agreement to develop TAK-999 (ARO-AAT), a Phase 2 investigational RNA interference (RNAi) therapy in development to treat alpha-1 antitrypsin-associated liver disease (AATLD). ARO-AAT is a potential first-in-class therapy designed to reduce the production of mutant alpha-1 antitrypsin protein, the cause of AATLD progression.
Beacon Discovery	U.S.	Collaboration agreement for the G-protein coupled receptor drug discovery and development program to identify drug candidates for a range of gastrointestinal disorders. The agreement grants Takeda worldwide rights to develop, manufacture and commercialize products resulting from the collaboration.
Cerevance	U.S.	Multi-year research alliance to identify novel target proteins expressed in the central nervous system and to develop new therapies against them for certain GI disorders. Goal of the collaboration is to select, confirm and validate targets from gene expression data sets generated by Cerevance’s NETSeq technology.
Cour Pharmaceutical Development Company	U.S.	Takeda has acquired an exclusive global license to develop and commercialize the investigational medicine TIMP-GLIA (TAK-101), an immune modifying nanoparticle containing gliadin proteins.
Engitix	U.K.	Collaboration and licensing agreement to utilize Engitix’s liver fibrosis platform to conduct research activities and to nominate, confirm, and validate potential targets against which Takeda may advance new therapeutic programs.
Enterome	France	Collaboration agreement to research and develop microbiome targets thought to play crucial roles in gastrointestinal disorders, including inflammatory bowel diseases (e.g. ulcerative colitis). The agreement includes a global license and co-development of EB8018/TAK-018 in Crohn’s disease.

Finch Therapeutics	U.S.	Global agreement to develop TAK-524, a live biotherapeutic product composed of cultured bacterial strains linked to favorable clinical outcomes in studies of microbiota transplantations in inflammatory bowel disease. Under the terms of the agreement, Takeda has the exclusive worldwide rights to develop and commercialize TAK-524 and rights to follow-on products in inflammatory bowel diseases. Following a contract amendment in Aug 2021, Takeda assumes sole responsibility for development of TAK-524, prior to the start of clinical development.
Genevant Sciences Corporation	U.S.	Collaboration and License Agreements to leverage Genevant’s hepatic stellate cell-partitioning LNP platform to deliver Takeda-designed RNAi oligonucleotides intended to halt or reverse the progression of liver fibrosis, and to deliver Takeda-designed non-viral gene therapies for the treatment of specified rare liver diseases.
Hemoshear Therapeutics	U.S.	Collaboration agreement for novel target and therapeutic development for liver diseases, using Hemoshear’s proprietary REVEAL-Tx drug discovery platform.
HiFiBio	U.S.	Collaboration agreement for functional therapeutics high-throughput antibody discovery platform that enables identification of antibodies for rare events for discovery of therapeutic antibodies for GI therapeutic areas.
NuBiyota	Canada	Agreement for the development of Microbial Ecosystem Therapeutic products for gastroenterology indications.
Mirum Pharmaceuticals <sup>‡</sup>	U.S.	Exclusive licensing agreement for the development and commercialization of maralixibat in Japan for Alagille syndrome (ALGS), progressive familial intrahepatic cholestasis (PFIC), and biliary atresia (BA).
Phathom Pharmaceuticals	U.S.	Takeda has granted a license to Phathom Pharmaceuticals for the development and exclusive commercialization rights to vonoprazan in the U.S., Europe and Canada in exchange for upfront cash and equity, as well as future cash milestones and royalties on net sales.
Sosei Heptares	UK	Collaboration and License agreement to leverage Sosei Heptares’s StaR® technology and structural biology expertise with GPCRs to enable structure based drug discovery to advance novel therapeutics for gastroenterology diseases.
Theravance Biopharma	U.S.	Global license, development and commercialization agreement for TAK-954, a selective 5-HT4 receptor agonist for motility disorders.
UCSD/Fortis Advisors	U.S.	Technology license for the development of oral budesonide formulation (TAK-721) for treatment of eosinophilic esophagitis.

<sup>‡</sup> Executed since April 1, 2021

• **Plasma Derived Therapies**

Partner	Country of incorporation	Subject
Halozyme	U.S.	Agreement for the in-license of Halozyme’s proprietary ENHANZE™ platform technology to increase dispersion and absorption of HyQvia. Ongoing development work for a U.S. pediatric indication to treat primary immunodeficiencies and a Phase 3 indication in Chronic Inflammatory Demyelinating Polyradiculoneuropathy.
Kamada	Israel	In-license agreement to develop and commercialize IV Alpha-1 proteinase inhibitor (Glassia); Exclusive supply and distribution of Glassia in the U.S., Canada, Australia and New Zealand; work on post market commitments ongoing.
ProThera Biologics	U.S.	Global licensing agreement to develop a novel plasma-derived Inter-alpha Inhibitor Proteins (IAIP) therapy for the treatment of acute inflammatory conditions.

• **Vaccines**

Partner	Country of incorporation	Subject
U.S. Government - The Biomedical Advanced Research and Development Authority (BARDA)	U.S.	Partnership to develop TAK-426, a Zika vaccine candidate, for the U.S. with the option to use data generated for filing also in affected regions around the world.
HilleVax, Inc. ‡	U.S.	Collaboration with Frazier Healthcare Partners to launch HilleVax, Inc., a biopharmaceutical company to advance the development and commercialization of norovirus vaccine candidate HIL-214 (formerly TAK-214). HilleVax has exclusive global development rights and commercialization rights worldwide outside of Japan in exchange for upfront consideration, as well as future cash milestones and royalties on net sales (Takeda retains commercialization rights in Japan).
Novavax	U.S.	Partnership for the development, manufacturing and commercialization of over 250 million doses per year of TAK-019 (NVX-CoV2373), Novavax' COVID-19 vaccine candidate, in Japan, which is being funded by the Government of Japan's Ministry of Health, Labour and Welfare (MHLW) and Agency for Medical Research and Development (AMED). Takeda finalized an agreement with the MHLW to supply 150 million doses of Novavax' vaccine candidate, subject to a number of factors including regulatory approval.
Moderna	U.S.	Three-way agreement with Moderna and the Government of Japan's Ministry of Health Labour & Welfare (MHLW) to import and distribute Moderna's COVID-19 vaccine, known as Spikevax Intermuscular Injection in Japan. The MHLW granted special approval for the primary series in May 2021 and regulatory approval for a 50 µg booster dose in December 2021. Takeda will deliver a total of 93 million doses (50 µg booster dose) to Japan in 2022, in addition to the 50 million doses (100 µg) delivered in 2021.

‡ Executed since April 1, 2021

• **Other / Multiple Therapeutic Area**

Partner	Country of incorporation	Subject
Bridge Medicines	U.S.	Partnership with Tri-Institutional Therapeutics Discovery Institute, Bay City Capital and Deerfield Management in the establishment of Bridge Medicines. Bridge Medicines will give financial, operational and managerial support to move projects seamlessly from a validating, proof-of-concept study to an in-human clinical trial.
Center for iPS Cell Research Application, Kyoto University (CiRA)	Japan	Collaboration agreement for clinical applications of iPS cells in Takeda strategic areas including applications in neurosciences, oncology and GI as well as discovery efforts in additional areas of compelling iPSC translational science.
Charles River Laboratories	U.S.	Collaboration on multiple integrated programs across Takeda's core therapeutic areas using Charles River Laboratories' end-to-end drug discovery and safety assessment platform to progress these programs towards candidate status.
Evotec SE	Germany	Research alliance to support Takeda's growing number of research stage gene therapy discovery programs. Evotec and Takeda have also entered into a multi-RNA target alliance to discover and develop RNA targeting small molecule therapeutics for targets that are difficult to address via more conventional approaches.
Massachusetts Institute of Technology	U.S.	MIT-Takeda Program to fuel the development and application of artificial intelligence (AI) capabilities to benefit human health and drug development. Centered within the Abdul Latif Jameel Clinic for Machine Learning in Health (J-Clinic), the new program will leverage the combined expertise of both organizations, and is supported by Takeda's three-year investment (with the potential for a two-year extension).
Portal Instruments	U.S.	Agreement for the development and commercialization of Portal's jet injector drug delivery device for potential use with Takeda's investigational or approved biologic medicines.
Schrödinger	U.S.	Agreement for the multi-target research collaboration combining Schrödinger's in silico platform-driven drug discovery capabilities with Takeda's deep therapeutic area knowledge and expertise in structural biology.
Stanford University	U.S.	Collaboration agreement with Stanford University to form the Stanford Alliance for Innovative Medicines to more effectively develop innovative treatments and therapies.
Tri-Institutional Therapeutics Discovery Institute (Tri-I TDI)	U.S.	Agreement for the collaboration of academic institutions and industry to more effectively develop innovative treatments and therapies.
Twist Bioscience	U.S.	Agreement and license for Takeda to access Twist's "Library of Libraries," a panel of synthetic antibody phage display libraries derived only from sequences that exist in the human body. Together, the companies will work to discover, validate and optimize new antibody candidates.



• **Completed Partnerships [Update since April 1st, 2021]**

Partner	Country of incorporation	Subject
CoVIg-19 Plasma Alliance	-	Alliance formed by Takeda and CSL Behring to develop a potential plasma-derived therapy for treating COVID-19. The alliance goal is the development of a non-branded hyperimmune globulin medicine (CoVIg-19) with the potential to treat hospitalized adult patients with COVID-19.
Maverick Therapeutics	U.S.	Collaboration agreement for the development of Maverick Therapeutics' T-cell engagement platform created specifically to improve the utility of T-cell redirection therapy for the treatment of cancer. Under the agreement, Takeda has the exclusive option to acquire Maverick Therapeutics 5 years after partnership initiation in 2017 which was exercised April 2021.
Samsung Bioepis	Korea	Strategic collaboration agreement to jointly fund and co-develop multiple novel biologic therapies in unmet disease areas. The program's first therapeutic candidate is TAK-671, which is intended to treat severe acute pancreatitis.
Silence Therapeutics	U.K.	Technology Evaluation Agreement with Silence Therapeutics to access their GalNAc-siRNA technology platform. The objective of the evaluation is to identify a GalNAc-conjugated siRNA that inhibits expression of a proprietary Takeda target.
Centre d'Immunologie de Marseille-Luminy	France	Collaboration agreement to bring together expertise and knowledge in innate biology with Takeda's BacTrap capabilities to identify novel targets and pathways in myeloid cells.
Shattuck Labs	U.S.	Collaboration agreement to explore and develop checkpoint fusion proteins utilizing Shattuck's unique Agonist Redirected Checkpoint (ARC) <sup>TM</sup> platform which enables combination immunotherapy with a single product. Takeda will have the option to take an exclusive license to further develop and commercialize TAK-252/SL-279252). Takeda and Shattuck Labs mutually agree to terminate Collaboration Agreement in November 2021.
Rani Therapeutics	U.S.	Research collaboration agreement to evaluate a micro tablet pill technology for oral delivery of FVIII therapy in hemophilia.
Biological E. Limited	India	Takeda agreed to transfer existing measles and acellular pertussis vaccine bulk production technology to develop low-cost combination vaccines for India, China and low- and middle-income countries.
Seattle Collaboration	U.S.	Agreement for SPRInT (Seattle Partnership for Research on Innovative Therapies) to accelerate the translation of Fred Hutchinson Cancer Research Center's and University of Washington's cutting-edge discoveries into treatments for human disease (focusing on Oncology, GI and Neuroscience).

■ **Clinical study protocol summaries**

Clinical study protocol summaries are disclosed on the English-language web-site (<https://clinicaltrials.takeda.com/>) and clinical study protocol information in the Japanese-language is disclosed on the Japanese-language web-site (<https://www.takeda.com/what-we-do/research-and-development/takeda-clinical-trial-transparency/>).

We anticipate that this disclosure will assure transparency of information on Takeda's clinical trials for the benefit of healthcare professionals, their patients and other stakeholders, which we believe will contribute to the appropriate use of Takeda's products worldwide.



## 2. Supplementary Financial Information

### Revenue by region

Year to date

(Bn JPY)	Reported *1				Underlying *1
	FY20Q3 YTD	FY21Q3 YTD	YOY		YOY
Total revenue	2,427.5	2,695.7	268.2	11.0%	7.1%
Japan *2	435.1	530.2	95.1	21.9%	6.9%
% of revenue	17.9%	19.7%	1.7pt		
United States	1,189.0	1,297.0	108.1	9.1%	4.5%
% of revenue	49.0%	48.1%	-0.9pt		
Europe and Canada	500.0	541.0	41.0	8.2%	9.7%
% of revenue	20.6%	20.1%	-0.5pt		
Growth and Emerging Markets *3	303.5	327.5	24.0	7.9%	14.2%
% of revenue	12.5%	12.1%	-0.4pt		
Asia (excluding Japan)	119.2	139.8	20.6	17.3%	21.9%
% of revenue	4.9%	5.2%	0.3pt		
Latin America	95.4	93.5	-1.9	-2.0%	16.0%
% of revenue	3.9%	3.5%	-0.5pt		
Russia/CIS	38.7	43.6	4.9	12.5%	8.5%
% of revenue	1.6%	1.6%	0.0pt		
Other *4	50.2	50.6	0.4	0.8%	0.7%
% of revenue	2.1%	1.9%	-0.2pt		
Of which royalty / service income *2	69.0	210.6	141.5	205.0%	

\*1 Revenue amount is classified into countries or regions based on the customer location.

\*2 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21Q3YTD.

\*3 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa.

\*4 Other region includes Middle East, Oceania and Africa.

**Quarterly**

<b>(Bn JPY)</b>	Reported *1											
	FY20				FY21							
	Q1	Q2	Q3	Q4	Q1	YOY	Q2	YOY	Q3	YOY	Q4	YOY
Total revenue	801.9	788.9	836.8	770.3	949.6	18.4%	844.8	7.1%	901.3	7.7%		
Japan *2	144.0	138.3	152.7	124.6	259.0	79.8%	131.9	-4.6%	139.4	-8.7%		
% of revenue	18.0%	17.5%	18.3%	16.2%	27.3%		15.6%		15.5%			
United States	402.6	383.5	402.8	379.0	412.2	2.4%	426.2	11.1%	458.6	13.9%		
% of revenue	50.2%	48.6%	48.1%	49.2%	43.4%		50.4 %		50.9 %			
Europe and Canada	157.6	169.6	172.8	166.2	178.7	13.4%	175.2	3.3%	187.0	8.2%		
% of revenue	19.6%	21.5%	20.7%	21.6%	18.8%		20.7 %		20.7 %			
Growth and Emerging Markets *3	97.6	97.5	108.4	100.5	99.7	2.1%	111.5	14.4%	116.3	7.3%		
% of revenue	12.2%	12.4%	13.0%	13.0%	10.5%		13.2 %		12.9 %			
Asia (excluding Japan)	36.9	41.4	40.9	37.1	40.3	9.3%	49.4	19.3%	50.1	22.4%		
% of revenue	4.6%	5.2%	4.9%	4.8%	4.2%		5.8 %		5.6 %			
Latin America	30.8	28.2	36.4	26.2	30.1	-2.3%	31.3	11.1%	32.2	-11.7%		
% of revenue	3.8%	3.6%	4.4%	3.4%	3.2%		3.7 %		3.6 %			
Russia/CIS	13.0	8.6	17.1	18.8	12.3	-5.4%	12.8	48.0%	18.5	8.4%		
% of revenue	1.6%	1.1%	2.0%	2.4%	1.3%		1.5 %		2.1 %			
Other *4	16.9	19.3	14.0	18.3	17.0	0.3%	18.0	-6.3%	15.5	11.1%		
% of revenue	2.1%	2.4%	1.7%	2.4%	1.8%		2.1 %		1.7 %			
Of which royalty / service income *2	18.1	28.2	22.8	23.4	157.7	773.2%	25.4	-9.8%	27.4	20.5%		

\*1 Revenue amount is classified into countries or regions based on the customer location.

\*2 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21Q1.

\*3 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa.

\*4 Other region includes Middle East, Oceania and Africa.

**Product Sales Analysis (vs PY Reported Actual)** (Sales amount includes royalty income and service income)

- Year to date

(Bn JPY)	FY20Q3 YTD	Reported		US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
		FY21Q3 YTD	YOY										
<b>GI</b>	<b>588.8</b>	<b>665.7</b>	<b>13.1%</b>	<b>373.5</b>	<b>8.0%</b>	<b>83.1</b>	<b>18.4%</b>	<b>146.0</b>	<b>24.4%</b>	<b>49.5</b>	<b>11.6%</b>	<b>13.5</b>	<b>22.9%</b>
ENTYVIO	319.3	395.4	23.8%	266.0	21.3%	8.6	30.6%	102.2	26.9%	18.6	43.3%		
TAKECAB-F *1	64.1	78.4	22.2%	—	-	73.3	16.3%	—	-	5.0	375.9%		
GATTEX/REVESTIVE	50.1	56.6	12.9%	46.4	8.1%	0.8	-	8.2	26.4%	1.3	71.8%		
DEXILANT	43.5	40.1	-7.6%	23.2	-15.1%	—	-	7.7	23.6%	9.2	-6.6%		
PANTOLOC/CONTROLOC*2	32.4	30.1	-7.1%	1.7	7.8%	—	-	19.7	14.5%	8.6	-36.3%		
LIALDA/MEZAVANT *3	18.7	19.0	1.7%	5.5	-28.5%							13.5	22.9%
PENTASA	17.8	15.8	-11.6%	15.8	-11.6%								
AMITIZA	18.8	5.9	-68.8%	5.3	-71.3%			—	-	0.6	76.1%		
RESOLOR/MOTTEGRITY	8.5	10.1	18.6%	7.6	25.2%	—	-	2.5	1.8%	—	-100.0%		
ALOFISEL	0.6	1.4	139.5%	—	-	—	-	1.1	151.0%	0.2	93.5%		
Others	14.9	13.0	-13.1%	2.0	-57.2%	0.5	-15.1%	4.6	13.9%	5.9	4.4%		
<b>Rare Diseases</b>	<b>446.7</b>	<b>462.9</b>	<b>3.6%</b>	<b>204.5</b>	<b>1.4%</b>	<b>23.1</b>	<b>0.8%</b>	<b>115.7</b>	<b>11.8%</b>	<b>80.1</b>	<b>0.4%</b>	<b>39.6</b>	<b>1.8%</b>
<b>Rare Metabolic</b>	<b>121.8</b>	<b>133.4</b>	<b>9.5%</b>	<b>28.2</b>	<b>7.3%</b>	<b>2.1</b>	<b>20.1%</b>	<b>36.0</b>	<b>12.8%</b>	<b>27.5</b>	<b>19.9%</b>	<b>39.6</b>	<b>1.8%</b>
ELAPRASE	51.5	57.7	12.0%	15.2	2.9%	1.0	18.8%	20.3	9.9%	21.2	21.6%		
REPLAGAL *3	38.9	39.6	1.8%	—	-							39.6	1.8%
VPRIV	28.9	32.2	11.4%	13.1	11.0%	1.1	21.4%	11.7	9.5%	6.3	14.5%		
NATPARA/NATPAR	2.5	3.9	56.9%	0.0	-	—	-	3.9	46.9%	0.0	0.3%		
<b>Rare Hematology</b>	<b>218.6</b>	<b>211.6</b>	<b>-3.2%</b>	<b>91.1</b>	<b>-1.2%</b>	<b>19.5</b>	<b>-3.0%</b>	<b>52.4</b>	<b>-0.4%</b>	<b>48.6</b>	<b>-9.4%</b>		
ADVATE	97.1	89.3	-8.0%	42.5	-6.9%	4.6	-9.5%	20.3	-14.8%	22.0	-2.9%		
ADYNOVATE/ADYNOVI	43.8	45.9	4.8%	19.7	0.7%	11.5	-2.1%	10.6	12.4%	4.1	34.7%		
FEIBA *4	34.2	29.0	-15.4%	8.9	10.7%	0.6	-19.8%	7.9	-0.8%	11.6	-33.8%		
RECOMBINATE	10.5	9.6	-8.3%	8.9	-3.6%	—	-	0.6	-18.2%	0.1	-84.3%		
HEMOFIL/IMMUNATE/IMMUNINE*4	13.2	13.5	2.3%	2.7	3.9%	—	-	3.5	-0.4%	7.3	3.1%		
Other PDT Products *4 *6	2.6	3.0	17.5%	0.0	-	—	-	2.7	22.4%	0.3	-11.8%		
Others *7	17.2	21.3	23.7%	8.4	17.1%	2.9	10.4%	6.8	38.2%	3.2	27.9%		
<b>Hereditary Angioedema</b>	<b>106.4</b>	<b>117.7</b>	<b>10.7%</b>	<b>84.9</b>	<b>2.2%</b>	<b>1.5</b>	<b>39.0%</b>	<b>27.4</b>	<b>43.7%</b>	<b>4.0</b>	<b>24.6%</b>		
TAKHZYRO	65.9	78.4	19.0%	60.7	7.7%	—	-	15.9	81.2%	1.9	136.2%		
FIRAZYR	20.1	21.5	6.8%	10.7	-1.8%	1.5	39.0%	7.5	23.4%	1.8	-13.5%		
CINRYZE *4	17.3	14.7	-14.8%	10.3	-18.5%	—	-	4.0	-4.8%	0.3	-3.3%		
KALBITOR	3.1	3.1	1.2%	3.1	1.2%	—	-	—	-	—	-100.0%		
<b>Others</b>	<b>—</b>	<b>0.2</b>	<b>-</b>	<b>0.2</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>		

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 generic name: pantoprazole

\*3 License-out product : Regional breakdown is not available due to contract.

\*4 PDT products

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*7 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRID and Other Hemophilia.

Table of Contents

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
	FY20Q3 YTD	FY21Q3 YTD	YOY										
<b>PDT Immunology</b>	<b>313.0</b>	<b>363.2</b>	<b>16.0%</b>	<b>238.9</b>	<b>11.1%</b>							<b>124.3</b>	<b>26.8%</b>
immunoglobulin *1	248.0	278.3	12.2%	207.4	10.1%							70.9	18.8%
albumin *1	43.6	61.5	41.0%	13.4	30.7%							48.1	44.2%
Others *1 *6	21.4	23.4	9.5%	18.1	10.3%							5.4	6.9%
<b>Oncology</b>	<b>318.5</b>	<b>359.1</b>	<b>12.8%</b>	<b>171.5</b>	<b>11.8%</b>	<b>67.1</b>	<b>8.0%</b>	<b>59.8</b>	<b>8.2%</b>	<b>53.8</b>	<b>33.9%</b>	<b>6.9</b>	<b>-7.8%</b>
VELCADE *2	75.9	84.5	11.3%	81.8	13.3%							2.7	-27.6%
LEUPLIN/ENANTONE	75.3	82.2	9.2%	18.1	87.9%	23.3	-20.1%	24.6	3.4%	16.2	27.6%		
NINLARO	67.9	70.7	4.2%	43.0	-4.0%	4.7	21.2%	10.4	2.2%	12.7	40.1%		
ADCETRIS	44.4	51.8	16.7%			8.8	3.5%	21.0	11.5%	22.0	28.9%		
ICLUSIG *2	26.3	26.7	1.6%	22.5	-0.1%							4.2	11.9%
VECTIBIX	18.4	19.4	5.3%			19.4	5.3%						
ALUNBRIG	6.5	10.1	56.2%	5.1	17.5%	0.8	-	2.8	80.7%	1.4	144.0%		
Others	3.9	13.7	247.4%	1.1	-	10.2	340.9%	1.0	12.2%	1.4	94.2%		
<b>Neuroscience</b>	<b>315.1</b>	<b>362.6</b>	<b>15.1%</b>	<b>281.3</b>	<b>16.1%</b>	<b>25.3</b>	<b>-15.7%</b>	<b>48.4</b>	<b>29.2%</b>	<b>7.6</b>	<b>45.3%</b>		
VYVANSE/ELVANSE	202.4	245.0	21.0%	202.2	17.7%	0.2	8,176.3%	35.6	38.3%	6.9	43.1%		
TRINTELLIX	52.7	63.0	19.6%	59.0	14.8%	4.1	216.7%			—	-100.0%		
INTUNIV	14.9	12.5	-15.7%	0.1	-81.9%	3.5	-55.7%	8.3	36.6%	0.6	79.6%		
ADDERALL XR	13.4	16.0	19.5%	14.4	18.7%	—	-	1.5	27.8%	—	-		
ROZEREM	9.5	9.4	-0.8%	0.1	-83.7%	9.3	2.4%	0.0	-	0.1	66.7%		
Others *7	22.3	16.7	-25.1%	5.6	-8.3%	8.2	-30.2%	2.9	-34.2%	0.0	-56.2%		
<b>Others *3</b>	<b>445.4</b>	<b>482.2</b>	<b>8.2%</b>										
AZILVA-F *4	62.8	60.1	-4.4%	—	-	60.1	-4.4%	—	-	—	-		
LOTRIGA	24.5	24.8	1.2%			24.8	1.2%						
AIPHAGAN	12.2	12.0	-1.9%	—	-	12.0	-1.9%	—	-	—	-		
FOSRENOL *2	10.2	10.2	-0.2%	1.3	-13.3%							8.9	2.0%
ACTOVEGIN	8.3	11.0	33.5%	—	-	—	-	0.6	48.9%	10.4	32.7%		

\*1 PDT products

\*2 License-out product : Regional breakdown is not available due to contract.

\*3 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21Q3YTD.

\*4 The figures include the amounts of fixed dose combinations.

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*7 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL

Table of Contents

- Quarterly
- Q1

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
	FY20 Q1	FY21 Q1	YOY										
<b>GI</b>	<b>186.9</b>	<b>210.5</b>	<b>12.6%</b>	<b>117.6</b>	<b>3.3%</b>	<b>25.6</b>	<b>15.9%</b>	<b>47.1</b>	<b>36.1%</b>	<b>16.0</b>	<b>23.6%</b>	<b>4.2</b>	<b>20.5%</b>
ENTYVIO	101.2	125.4	23.9%	83.7	17.1%	2.5	26.6%	32.7	35.6%	6.4	78.9%		
TAKECAB-F *1	20.2	24.3	20.1%	—	-	22.9	15.0%	—	-	1.4	346.2%		
GATTEX/REVESTIVE	17.5	18.1	3.7%	15.2	-1.1%	—	-	2.7	43.5%	0.2	3.6%		
DEXILANT	13.6	10.8	-20.7%	6.0	-31.3%	—	-	2.2	18.3%	2.5	-14.3%		
PANTOLOC/CONTROLOC*2	9.2	10.4	13.8%	0.7	39.8%	—	-	6.7	37.4%	3.1	-19.1%		
LIALDA/MEZAVANT *3	5.5	6.4	16.2%	2.2	8.6%							4.2	20.5%
PENTASA	6.2	4.8	-21.6%	4.8	-21.6%								
AMITIZA	6.3	2.1	-65.8%	2.0	-67.8%			—	-	0.1	133.0%		
RESOLOR/MOTTEGRITY	2.7	3.2	16.9%	2.2	10.2%	—	-	1.0	43.4%	—	-100.0%		
ALOFISEL	0.0	0.4	3,556.0%	—	-	—	-	0.3	4,796.0%	0.1	1,513.3%		
Others	4.5	4.5	0.0%	0.7	-41.8%	0.2	-7.1%	1.5	26.6%	2.1	10.8%		
<b>Rare Diseases</b>	<b>155.0</b>	<b>155.5</b>	<b>0.3%</b>	<b>71.2</b>	<b>-3.9%</b>	<b>7.5</b>	<b>-2.4%</b>	<b>38.6</b>	<b>11.8%</b>	<b>24.1</b>	<b>-9.0%</b>	<b>14.1</b>	<b>15.2%</b>
<b>Rare Metabolic</b>	<b>39.9</b>	<b>44.3</b>	<b>10.9%</b>	<b>9.4</b>	<b>5.5%</b>	<b>0.7</b>	<b>-2.6%</b>	<b>11.8</b>	<b>17.1%</b>	<b>8.3</b>	<b>3.7%</b>	<b>14.1</b>	<b>15.2%</b>
ELAPRASE	17.6	18.6	5.5%	5.0	0.1%	0.4	-5.1%	6.7	13.9%	6.4	2.5%		
REPLAGAL *3	12.2	14.1	15.2%	—	-							14.1	15.2%
VPRIV	9.3	10.5	11.9%	4.4	13.8%	0.3	0.8%	3.9	12.7%	1.8	7.9%		
NATPARA/NATPAR	0.7	1.2	56.8%	-0.0	-	—	-	1.2	64.6%	0.0	45.1%		
<b>Rare Hematology</b>	<b>76.8</b>	<b>72.2</b>	<b>-5.9%</b>	<b>33.3</b>	<b>-0.4%</b>	<b>6.4</b>	<b>-2.6%</b>	<b>18.0</b>	<b>-6.0%</b>	<b>14.6</b>	<b>-17.6%</b>		
ADVATE	33.7	30.7	-8.9%	15.1	-11.1%	1.6	-5.4%	7.1	-13.0%	6.9	0.6%		
ADYNOVATE/ADYNOVI	15.3	15.4	0.6%	6.8	-5.5%	3.7	-3.4%	3.6	6.1%	1.2	50.6%		
FEIBA *4	12.9	11.4	-11.3%	3.9	60.3%	0.2	-8.8%	3.2	-2.3%	4.1	-40.7%		
RECOMBINATE	3.7	3.7	-0.9%	3.5	4.9%	—	-	0.2	-8.0%	0.0	-91.1%		
HEMOFIL/IMMUNATE/IMMUNINE*4	4.4	3.3	-25.6%	0.9	12.8%	—	-	1.0	-36.7%	1.4	-31.4%		
Other PDT Products *4 *6	0.9	0.9	-1.1%	0.0	-	—	-	0.8	15.6%	0.0	-81.4%		
Others *7	5.9	6.9	16.4%	3.1	15.1%	0.9	8.5%	2.1	16.3%	0.8	32.8%		
<b>Hereditary Angioedema</b>	<b>38.3</b>	<b>39.0</b>	<b>1.8%</b>	<b>28.5</b>	<b>-10.1%</b>	<b>0.4</b>	<b>2.9%</b>	<b>8.9</b>	<b>64.8%</b>	<b>1.3</b>	<b>48.2%</b>		
TAKHZYRO	23.2	25.5	9.6%	19.9	-5.7%	—	-	4.9	140.6%	0.6	572.2%		
FIRAZYR	8.1	6.9	-15.1%	3.4	-34.7%	0.4	2.9%	2.6	36.1%	0.5	-15.7%		
CINRYZE *4	5.9	5.6	-5.7%	4.1	-4.9%	—	-	1.3	-5.8%	0.1	-26.9%		
KALBITOR	1.1	1.1	2.8%	1.1	2.8%	—	-	—	-	—	-		
<b>Others</b>	<b>—</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>		

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 generic name: pantoprazole

\*3 License-out product : Regional breakdown is not available due to contract.

\*4 PDT products

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*7 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRID and Other Hemophilia.

■ Q1

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
	FY20 Q1	FY21 Q1	YOY										
<b>PDT Immunology</b>	<b>105.3</b>	<b>107.2</b>	<b>1.8%</b>	<b>70.3</b>	<b>-5.4%</b>							<b>36.9</b>	<b>19.2%</b>
immunoglobulin *1	85.1	81.6	-4.1%	59.0	-10.7%							22.6	19.0%
albumin *1	13.0	17.8	36.8%	5.3	100.9%							12.5	20.6%
Others *1 *6	7.2	7.8	9.1%	6.0	7.8%							1.8	13.5%
<b>Oncology</b>	<b>108.0</b>	<b>121.4</b>	<b>12.4%</b>	<b>60.4</b>	<b>20.7%</b>	<b>21.0</b>	<b>-11.0%</b>	<b>21.2</b>	<b>15.4%</b>	<b>16.6</b>	<b>23.7%</b>	<b>2.1</b>	<b>-15.9%</b>
VELCADE *2	24.2	30.1	24.6%	29.4	27.3%							0.8	-30.8%
LEUPLIN/ENANTONE	27.4	26.2	-4.3%	4.8	128.5%	7.5	-41.4%	9.7	17.7%	4.2	-1.7%		
NINLARO	22.9	24.4	6.3%	15.4	-1.3%	1.5	19.3%	3.5	3.5%	4.1	46.0%		
ADCETRIS	15.1	17.2	14.2%			2.8	-2.4%	7.0	13.2%	7.5	22.9%		
ICLUSIG *2	9.2	10.4	12.3%	9.1	15.0%							1.3	-3.6%
VECTIBIX	6.2	6.2	0.1%			6.2	0.1%						
ALUNBRIG	2.0	3.1	54.4%	1.7	18.0%	0.2	-	0.8	89.6%	0.4	160.4%		
Others	0.9	3.8	300.0%	0.1	-	2.8	418.7%	0.3	25.9%	0.5	231.1%		
<b>Neuroscience</b>	<b>106.9</b>	<b>113.4</b>	<b>6.1%</b>	<b>87.3</b>	<b>8.7%</b>	<b>7.5</b>	<b>-39.9%</b>	<b>15.9</b>	<b>37.0%</b>	<b>2.8</b>	<b>10.1%</b>		
VYVANSE/ELVANSE	66.0	79.2	20.0%	65.2	16.6%	0.0	-	11.5	48.3%	2.5	7.8%		
TRINTELLIX	16.9	17.9	5.9%	16.7	0.9%	1.1	273.0%			—	-		
INTUNIV	5.6	3.3	-42.5%	-0.0	-	0.4	-89.1%	2.7	44.6%	0.2	59.1%		
ADDERALL XR	5.3	3.9	-24.9%	3.5	-27.5%	—	-	0.4	4.0%	—	-		
ROZEREM	3.0	3.2	6.9%	0.1	485.1%	3.1	4.1%	—	-	0.0	8.5%		
Others *7	10.0	5.9	-41.2%	1.8	-29.7%	2.9	-51.1%	1.2	-21.7%	0.0	-51.2%		
<b>Others *3</b>	<b>139.8</b>	<b>241.6</b>	<b>72.8%</b>										
AZILVA-F *4	20.9	22.6	8.6%	—	-	22.6	8.6%	—	-	—	-		
LOTRIGA	8.1	7.8	-3.0%			7.8	-3.0%						
AIPHAGAN	4.0	4.6	15.0%	—	-	4.6	15.0%	—	-	—	-		
FOSRENOL *2	3.2	3.4	4.7%	0.5	-30.2%							2.8	15.9%
ACTOVEGIN	1.7	3.2	87.2%	—	-	—	-	0.2	222.9%	3.0	81.8%		

\*1 PDT products

\*2 License-out product : Regional breakdown is not available due to contract.

\*3 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21Q1.

\*4 The figures include the amounts of fixed dose combinations.

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*7 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL

## ■ Q2

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
	FY20 Q2	FY21 Q2	YOY										
<b>GI</b>	<b>192.9</b>	<b>218.6</b>	<b>13.3%</b>	<b>123.2</b>	<b>11.3%</b>	<b>26.3</b>	<b>21.9%</b>	<b>48.0</b>	<b>16.7%</b>	<b>17.2</b>	<b>8.3%</b>	<b>3.9</b>	<b>6.5%</b>
ENTYVIO	105.7	130.5	23.4%	87.6	22.3%	2.9	42.1%	33.9	23.7%	6.2	31.5%		
TAKECAB-F *1	19.7	24.8	25.9%	—	-	23.1	19.2%	—	-	1.7	413.4%		
GATTEX/REVESTIVE	15.7	18.7	18.8%	15.4	16.1%	0.1	-	2.6	14.3%	0.6	176.0%		
DEXILANT	14.8	14.9	0.8%	8.7	-4.1%	—	-	2.6	11.7%	3.6	6.6%		
PANTOLOC/CONTROLOC*2	12.3	9.4	-23.4%	0.4	-22.4%	—	-	6.1	-5.9%	2.9	-45.2%		
LIALDA/MEZAVANT *3	6.1	5.3	-13.1%	1.4	-42.9%							3.9	6.5%
PENTASA	5.5	5.2	-5.6%	5.2	-5.6%								
AMITIZA	6.2	1.8	-71.6%	1.5	-75.5%			—	-	0.3	92.2%		
RESOLOR/MOTTEGRITY	2.2	3.2	43.3%	2.4	68.4%	—	-	0.8	1.8%	—	-100.0%		
ALOFISEL	0.3	0.4	52.2%	—	-	—	-	0.3	43.9%	0.1	93.5%		
Others	4.3	4.3	-0.0%	0.7	-13.9%	0.2	-0.4%	1.6	1.9%	1.9	4.4%		
<b>Rare Diseases</b>	<b>140.4</b>	<b>144.6</b>	<b>3.0%</b>	<b>62.1</b>	<b>-2.1%</b>	<b>6.6</b>	<b>-17.3%</b>	<b>37.8</b>	<b>9.2%</b>	<b>26.2</b>	<b>21.4%</b>	<b>11.9</b>	<b>-7.0%</b>
<b>Rare Metabolic</b>	<b>39.7</b>	<b>40.0</b>	<b>0.6%</b>	<b>9.1</b>	<b>1.6%</b>	<b>0.3</b>	<b>-62.8%</b>	<b>12.0</b>	<b>12.1%</b>	<b>6.7</b>	<b>1.9%</b>	<b>11.9</b>	<b>-7.0%</b>
ELAPRASE	16.7	16.2	-2.8%	4.9	-6.0%	-0.1	-	6.7	7.4%	4.6	-4.5%		
REPLAGAL *3	12.8	11.9	-7.0%	—	-							11.9	-7.0%
VPRIV	9.5	10.5	11.0%	4.2	6.7%	0.3	-9.8%	4.0	13.5%	2.1	19.7%		
NATPARA/NATPAR	0.8	1.3	72.1%	0.0	-	—	-	1.3	39.0%	0.0	7.6%		
<b>Rare Hematology</b>	<b>66.1</b>	<b>69.4</b>	<b>5.0%</b>	<b>28.1</b>	<b>-0.1%</b>	<b>6.0</b>	<b>-13.0%</b>	<b>17.1</b>	<b>-0.9%</b>	<b>18.2</b>	<b>31.9%</b>		
ADVATE	29.8	30.6	2.9%	13.3	-4.0%	1.4	-13.7%	7.1	-13.9%	8.8	46.9%		
ADYNOVATE/ADYNOVI	14.2	14.6	2.6%	6.5	11.4%	3.6	-12.5%	3.2	-5.3%	1.4	41.0%		
FEIBA *4	7.7	8.8	13.7%	2.2	-13.5%	0.2	-23.3%	2.8	39.2%	3.6	22.7%		
RECOMBINATE	3.2	2.6	-18.5%	2.4	-13.2%	—	-	0.2	-39.5%	0.0	-76.2%		
HEMOFIL/IMMUNATE/IMMUNINE*4	4.9	5.1	3.0%	0.8	-19.5%	—	-	1.3	32.8%	3.0	1.2%		
Other PDT Products *4 *6	0.8	1.1	31.3%	0.0	-	—	-	0.9	19.5%	0.2	124.3%		
Others *7	5.4	6.6	22.5%	2.8	38.2%	0.8	-11.7%	1.7	2.3%	1.3	62.7%		
<b>Hereditary Angioedema</b>	<b>34.6</b>	<b>35.2</b>	<b>1.8%</b>	<b>24.9</b>	<b>-5.4%</b>	<b>0.4</b>	<b>-12.9%</b>	<b>8.7</b>	<b>30.3%</b>	<b>1.3</b>	<b>7.2%</b>		
TAKHZYRO	20.5	22.1	7.6%	16.6	-2.2%	—	-	5.0	56.7%	0.4	36.7%		
FIRAZYR	7.1	7.5	5.9%	4.1	7.1%	0.4	-12.9%	2.4	14.7%	0.6	-13.8%		
CINRYZE *4	6.1	4.6	-24.3%	3.1	-31.9%	—	-	1.3	-6.5%	0.2	54.2%		
KALBITOR	0.9	1.1	14.0%	1.1	14.0%	—	-	—	-	—	-		
<b>Others</b>	<b>—</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>		

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 generic name: pantoprazole

\*3 License-out product : Regional breakdown is not available due to contract.

\*4 PDT products

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*7 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRID and Other Hemophilia.

■ Q2

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*4	YOY	Ex-US	YOY
	FY20 Q2	FY21 Q2	YOY										
<b>PDT Immunology</b>	<b>100.6</b>	<b>130.8</b>	<b>30.0%</b>	<b>86.8</b>	<b>28.9%</b>							<b>44.0</b>	<b>32.3%</b>
immunoglobulin *1	77.6	99.7	28.6%	76.5	32.1%							23.2	18.1%
albumin *1	15.6	24.0	53.8%	4.7	28.0%							19.3	61.7%
Others *1 *5	7.5	7.1	-4.6%	5.7	-2.8%							1.5	-11.2%
<b>Oncology</b>	<b>102.1</b>	<b>112.3</b>	<b>10.0%</b>	<b>51.0</b>	<b>0.8%</b>	<b>21.7</b>	<b>21.8%</b>	<b>19.1</b>	<b>4.6%</b>	<b>18.2</b>	<b>40.2%</b>	<b>2.2</b>	<b>-4.4%</b>
VELCADE *2	25.8	25.0	-3.3%	24.1	-1.8%							0.9	-31.7%
LEUPLIN/ENANTONE	22.5	27.6	23.0%	7.0	85.4%	7.6	1.5%	7.2	-0.9%	5.9	47.5%		
NINLARO	21.4	21.4	0.0%	12.0	-15.4%	1.5	25.6%	3.5	11.1%	4.5	53.4%		
ADCETRIS	15.5	16.9	9.3%			2.9	3.5%	7.2	1.9%	6.8	21.5%		
ICLUSIG *2	7.6	7.5	-1.6%	6.2	-6.6%							1.3	30.9%
VECTIBIX	5.7	6.6	15.0%			6.6	15.0%						
ALUNBRIG	2.3	3.1	38.8%	1.4	-8.0%	0.3	-	0.9	72.7%	0.5	162.1%		
Others	1.3	4.1	225.6%	0.4	-	2.9	315.5%	0.4	11.4%	0.5	95.0%		
<b>Neuroscience</b>	<b>100.9</b>	<b>120.3</b>	<b>19.2%</b>	<b>94.6</b>	<b>18.4%</b>	<b>8.5</b>	<b>21.1%</b>	<b>14.9</b>	<b>14.6%</b>	<b>2.3</b>	<b>117.6%</b>		
VYVANSE/ELVANSE	66.6	80.1	20.2%	66.7	16.8%	0.2	-	11.0	29.0%	2.1	126.2%		
TRINTELLIX	18.1	22.2	22.7%	20.9	17.7%	1.3	289.6%				-100.0%		
INTUNIV	3.3	4.2	26.6%	0.0	-78.6%	1.4	44.8%	2.6	23.0%	0.2	66.1%		
ADDERALL XR	3.7	5.7	52.9%	5.2	54.8%		-	0.5	35.9%		-		
ROZEREM	2.9	3.1	5.7%	0.1	-13.4%	3.0	5.8%	0.0	-	0.0	91.5%		
Others *6	6.3	5.1	-19.3%	1.7	21.4%	2.6	-11.0%	0.8	-60.7%		-100.0%		
<b>Others</b>	<b>152.0</b>	<b>118.2</b>	<b>-22.3%</b>										
AZILVA-F *3	19.1	17.7	-7.2%		-	17.7	-7.2%		-		-		
LOTRIGA	7.6	8.2	8.5%			8.2	8.5%						
AIPHAGAN	3.7	3.8	3.3%		-	3.8	3.3%		-		-		
FOSRENOL *2	3.3	3.6	9.9%	0.7	479.3%							2.9	-9.1%
ACTOVEGIN	3.2	3.5	8.8%		-		-	0.2	20.0%	3.3	8.1%		

\*1 PDT products

\*2 License-out product : Regional breakdown is not available due to contract.

\*3 The figures include the amounts of fixed dose combinations.

\*4 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*5 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*6 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL



■ Q3

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*5	YOY	Ex-US	YOY
	FY20 Q3	FY21 Q3	YOY										
<b>GI</b>	<b>209.0</b>	<b>236.6</b>	<b>13.2%</b>	<b>132.8</b>	<b>9.3%</b>	<b>31.2</b>	<b>17.8 %</b>	<b>51.0</b>	<b>22.3 %</b>	<b>16.3</b>	<b>5.0%</b>	<b>5.3</b>	<b>41.0%</b>
ENTYVIO	112.3	139.5	24.2%	94.7	24.4%	3.1	24.6 %	35.6	22.8 %	6.0	28.0%		
TAKECAB-F *1	24.2	29.3	21.0%	—	-	27.3	15.0 %	—	-	1.9	367.7%		
GATTEX/REVESTIVE	16.9	19.8	17.0%	15.8	10.6%	0.6	-	2.9	24.8 %	0.5	53.0%		
DEXILANT	15.1	14.4	-4.1%	8.5	-10.7%	—	-	2.9	42.2 %	3.1	-12.8%		
PANTOLOC/CONTROLOC*2	10.9	10.2	-6.5%	0.6	11.4%	—	-	6.9	18.0 %	2.7	-40.7%		
LIALDA/MEZAVANT *3	7.1	7.3	3.1%	2.0	-40.3%							5.3	41.0%
PENTASA	6.2	5.7	-6.9%	5.7	-6.9%								
AMITIZA	6.4	2.0	-69.2%	1.8	-70.8%			—	-	0.1	22.6%		
RESOLOR/MOTEGRITY	3.6	3.7	4.5%	3.0	13.7%	—	-	0.7	-27.5 %	—	-100.0%		
ALOFISEL	0.3	0.6	94.9%	—	-	—	-	0.5	121.3 %	0.1	5.6%		
Others	6.1	4.1	-32.0%	0.6	-77.0%	0.2	-31.3 %	1.5	16.6 %	1.9	-2.1%		
<b>Rare Diseases</b>	<b>151.3</b>	<b>162.8</b>	<b>7.6%</b>	<b>71.2</b>	<b>10.9%</b>	<b>9.0</b>	<b>24.0 %</b>	<b>39.3</b>	<b>14.5 %</b>	<b>29.7</b>	<b>-6.1%</b>	<b>13.6</b>	<b>-2.0%</b>
<b>Rare Metabolic</b>	<b>42.2</b>	<b>49.2</b>	<b>16.6%</b>	<b>9.8</b>	<b>15.0%</b>	<b>1.1</b>	<b>264.6 %</b>	<b>12.2</b>	<b>9.6 %</b>	<b>12.5</b>	<b>49.5%</b>	<b>13.6</b>	<b>-2.0%</b>
ELAPRASE	17.2	22.9	33.0%	5.2	16.3%	0.7	774.6 %	6.9	8.6 %	10.1	60.9%		
REPLAGAL *3	13.9	13.6	-2.0%	—	-							13.6	-2.0%
VPRIV	10.0	11.2	11.4%	4.5	12.6%	0.4	96.7 %	3.9	2.8 %	2.4	15.7%		
NATPARA/NATPAR	1.0	1.4	45.1%	0.0	-	—	-	1.4	41.5 %	0.0	-45.6%		
<b>Rare Hematology</b>	<b>75.8</b>	<b>70.0</b>	<b>-7.6%</b>	<b>29.8</b>	<b>-3.2%</b>	<b>7.1</b>	<b>7.1 %</b>	<b>17.3</b>	<b>6.7 %</b>	<b>15.8</b>	<b>-28.7%</b>		
ADVATE	33.7	28.0	-16.8%	14.1	-4.8%	1.6	-9.4 %	6.1	-17.7 %	6.3	-35.6%		
ADYNOVATE/ADYNOVI	14.3	15.9	11.5%	6.4	-2.1%	4.2	10.6 %	3.9	41.7 %	1.5	19.0%		
FEIBA *4	13.7	8.8	-35.6%	2.8	-8.1%	0.2	-27.9 %	1.9	-29.0 %	3.9	-49.2%		
RECOMBINATE	3.5	3.3	-7.0%	3.1	-4.1%	—	-	0.2	2.4 %	0.0	-83.0%		
HEMOFIL/IMMUNATE/IMMUNINE*4	3.9	5.2	33.5%	1.0	26.5%	—	-	1.2	27.6 %	3.0	38.7%		
Other PDT Products *4 *6	0.9	1.1	23.0%	-0.0	-4,573.1 %	—	-	1.0	31.8 %	0.1	-24.2%		
Others *7	5.8	7.7	32.2%	2.5	2.0%	1.2	35.8 %	3.0	106.2 %	1.0	-1.5%		
<b>Hereditary Angioedema</b>	<b>33.4</b>	<b>43.5</b>	<b>30.1%</b>	<b>31.4</b>	<b>26.0%</b>	<b>0.8</b>	<b>140.1 %</b>	<b>9.8</b>	<b>40.2 %</b>	<b>1.4</b>	<b>25.0%</b>		
TAKHZYRO	22.1	30.9	39.5%	24.2	32.5%	—	-	5.9	68.7 %	0.8	110.8%		
FIRAZYR	5.0	7.1	43.9%	3.2	74.9%	0.8	140.1 %	2.5	20.5 %	0.6	-11.3%		
CINRYZE *4	5.2	4.5	-14.2%	3.1	-17.8%	—	-	1.4	-2.1 %	0.0	-54.7%		
KALBITOR	1.1	1.0	-11.5%	1.0	-11.4%	—	-	—	-	—	-100.0%		
<b>Others</b>	<b>—</b>	<b>0.2</b>	<b>-</b>	<b>0.2</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>	<b>—</b>	<b>-</b>		

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 generic name: pantoprazole

\*3 License-out product : Regional breakdown is not available due to contract.

\*4 PDT products

\*5 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*6 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*7 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRID and Other Hemophilia.

■ Q3

(Bn JPY)	Reported			US	YOY	Japan	YOY	EUCAN	YOY	GEM*4	YOY	Ex-US	YOY
	FY20 Q3	FY21 Q3	YOY										
<b>PDT Immunology</b>	<b>107.1</b>	<b>125.2</b>	<b>16.9%</b>	<b>81.8</b>	<b>11.6%</b>							<b>43.4</b>	<b>28.4%</b>
immunoglobulin *1	85.4	97.0	13.6%	71.9	11.8%							25.1	19.2%
albumin *1	15.0	19.7	31.4%	3.5	-13.3%							16.3	47.5%
Others *1 *5	6.7	8.5	25.7%	6.4	28.3%							2.1	18.1%
<b>Oncology</b>	<b>108.4</b>	<b>125.4</b>	<b>15.7%</b>	<b>60.0</b>	<b>13.8%</b>	<b>24.3</b>	<b>17.7%</b>	<b>19.4</b>	<b>4.7%</b>	<b>18.9</b>	<b>38.0%</b>	<b>2.6</b>	<b>-3.3%</b>
VELCADE *2	25.9	29.3	13.4%	28.3	15.2%							1.0	-20.8%
LEUPLIN/ENANTONE	25.4	28.4	11.7%	6.3	67.7%	8.2	-7.6%	7.8	-6.9%	6.1	38.3%		
NINLARO	23.5	24.9	6.1%	15.6	3.9%	1.7	19.2%	3.5	-6.4%	4.2	23.8%		
ADCETRIS	13.8	17.6	27.7%			3.1	9.4%	6.9	21.9%	7.7	43.5%		
ICLUSIG *2	9.4	8.8	-6.2%	7.2	-9.6%							1.6	13.0%
VECTIBIX	6.5	6.6	1.7%			6.6	1.7%						
ALUNBRIG	2.2	3.9	75.6%	2.0	44.5%	0.3	-	1.0	81.6%	0.5	118.9%		
Others	1.7	5.8	234.8%	0.6	-	4.5	317.8%	0.3	2.5%	0.5	33.1%		
<b>Neuroscience</b>	<b>107.3</b>	<b>128.9</b>	<b>20.1%</b>	<b>99.5</b>	<b>20.9%</b>	<b>9.3</b>	<b>-11.3%</b>	<b>17.6</b>	<b>36.7%</b>	<b>2.5</b>	<b>51.8%</b>		
VYVANSE/ELVANSE	69.8	85.7	22.8%	70.3	19.6%	0.0	84.3%	13.1	38.4%	2.3	46.9%		
TRINTELLIX	17.7	23.0	29.6%	21.4	25.1%	1.6	151.5%			-	-		
INTUNIV	5.9	5.0	-13.9%	0.1	39.2%	1.7	-52.3%	3.0	42.9%	0.2	122.1%		
ADDERALL XR	4.4	6.3	44.4%	5.7	44.3%	-	-	0.6	45.7%	-	-		
ROZEREM	3.6	3.1	-12.6%	-0.1	-	3.2	-2.0%	-	-	0.0	184.0%		
Others *6	6.0	5.7	-4.0%	2.0	-1.8%	2.8	-7.3%	0.9	1.9%	-	-100.0%		
<b>Others</b>	<b>153.5</b>	<b>122.3</b>	<b>-20.3%</b>										
AZILVA-F *3	22.9	19.7	-13.8%	-	-	19.7	-13.8%	-	-	-	-		
LOTRIGA	8.8	8.7	-1.3%			8.7	-1.3%						
AIPHAGAN	4.6	3.6	-20.7%	-	-	3.6	-20.7%	-	-	-	-		
FOSRENOL *2	3.7	3.2	-13.6%	-0.0	-							3.2	2.4%
ACTOVEGIN	3.4	4.3	29.4%	-	-	-	-	0.2	11.3%	4.2	30.3%		

\*1 PDT products

\*2 License-out product : Regional breakdown is not available due to contract.

\*3 The figures include the amounts of fixed dose combinations.

\*4 GEM: Growth and Emerging Markets, which include Asia (excluding Japan), Latin America, Russia/CIS, Middle East, Oceania and Africa

\*5 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*6 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL

**Product Sales Analysis (Reported & Underlying Growth)**

(Bn JPY)	FY20 Reported				FY21 Reported & Underlying Growth															
	Q1	Q2	Q3	Q4	YOY			YOY				YOY				YOY				
					Q1	Reported	Underlying	Q2	Reported	Underlying	YTD Underlying	Q3	Reported	Underlying	YTD Underlying	Q4	Reported	Underlying	YTD Underlying	
<b>GI</b>	<b>186.9</b>	<b>192.9</b>	<b>209.0</b>	<b>189.0</b>	<b>210.5</b>	<b>12.6%</b>	<b>7.9%</b>	<b>218.6</b>	<b>13.3%</b>	<b>8.7%</b>	<b>8.3%</b>	<b>236.6</b>	<b>13.2%</b>	<b>6.2%</b>	<b>7.6%</b>					
ENTYVIO	101.2	105.7	112.3	110.0	125.4	23.9%	18.2%	130.5	23.4%	18.1%	18.1%	139.5	24.2%	15.5%	17.2%					
TAKECAB-F *1	20.2	19.7	24.2	20.7	24.3	20.1%	19.5%	24.8	25.9%	25.2%	22.3%	29.3	21.0%	20.2%	21.5%					
GATTEX/REVESTIVE	17.5	15.7	16.9	14.4	18.1	3.7%	0.3%	18.7	18.8%	14.1%	6.9%	19.8	17.0%	9.0%	7.6%					
DEXILANT	13.6	14.8	15.1	12.1	10.8	-20.7%	-24.6%	14.9	0.8%	-4.8%	-14.3%	14.4	-4.1%	-11.4%	-13.3%					
PANTOLOC/CONTROLOC*2	9.2	12.3	10.9	10.7	10.4	13.8%	4.4%	9.4	-23.4%	-28.1%	-14.0%	10.2	-6.5%	-12.5%	-13.5%					
LIALDA/MEZAVANT	5.5	6.1	7.1	6.8	6.4	16.2%	7.0%	5.3	-13.1%	-18.5%	-6.3%	7.3	3.1%	-5.3%	-5.9%					
PENTASA	6.2	5.5	6.2	5.3	4.8	-21.6%	-23.2%	5.2	-5.6%	-9.0%	-16.4%	5.7	-6.9%	-13.6%	-15.4%					
AMITIZA	6.3	6.2	6.4	2.4	2.1	-65.8%	-66.5%	1.8	-71.6%	-72.7%	-69.6%	2.0	-69.2%	-71.5%	-70.3%					
RESOLOR/MOTTEGRITY	2.7	2.2	3.6	2.7	3.2	16.9%	11.4%	3.2	43.3%	37.0%	22.8%	3.7	4.5%	-3.0%	12.0%					
ALOFISEL	0.0	0.3	0.3	0.2	0.4	3,556.0%	3,222.0%	0.4	52.2%	42.8%	166.8%	0.6	94.9%	80.9%	123.3%					
Others	4.5	4.3	6.1	3.7	4.5	0.0%	-6.2%	4.3	-0.0%	-4.2%	-5.2%	4.1	-32.0%	-35.5%	-17.6%					
<b>Rare Diseases</b>	<b>155.0</b>	<b>140.4</b>	<b>151.3</b>	<b>145.0</b>	<b>155.5</b>	<b>0.3%</b>	<b>-3.5%</b>	<b>144.6</b>	<b>3.0%</b>	<b>-0.7%</b>	<b>-2.2%</b>	<b>162.8</b>	<b>7.6%</b>	<b>1.3%</b>	<b>-1.0%</b>					
<b>Rare Metabolic</b>	<b>39.9</b>	<b>39.7</b>	<b>42.2</b>	<b>40.8</b>	<b>44.3</b>	<b>10.9%</b>	<b>6.6%</b>	<b>40.0</b>	<b>0.6%</b>	<b>-2.5%</b>	<b>2.1%</b>	<b>49.2</b>	<b>16.6%</b>	<b>11.1%</b>	<b>5.2%</b>					
ELAPRASE	17.6	16.7	17.2	17.3	18.6	5.5%	2.5%	16.2	-2.8%	-5.8%	-1.5%	22.9	33.0%	26.2%	7.8%					
REPLAGAL	12.2	12.8	13.9	12.9	14.1	15.2%	10.2%	11.9	-7.0%	-9.6%	0.2%	13.6	-2.0%	-5.2%	-1.8%					
VPRIV	9.3	9.5	10.0	9.7	10.5	11.9%	6.9%	10.5	11.0%	7.6%	7.2%	11.2	11.4%	5.6%	6.7%					
NATPARA/NATPAR	0.7	0.8	1.0	1.0	1.2	56.8%	39.1%	1.3	72.1%	61.5%	50.4%	1.4	45.1%	37.1%	45.2%					
<b>Rare Hematology</b>	<b>76.8</b>	<b>66.1</b>	<b>75.8</b>	<b>71.2</b>	<b>72.2</b>	<b>-5.9%</b>	<b>-9.6%</b>	<b>69.4</b>	<b>5.0%</b>	<b>1.1%</b>	<b>-4.6%</b>	<b>70.0</b>	<b>-7.6%</b>	<b>-13.2%</b>	<b>-7.6%</b>					
ADVATE	33.7	29.8	33.7	31.4	30.7	-8.9%	-12.9%	30.6	2.9%	-1.5%	-7.6%	28.0	-16.8%	-22.2%	-12.7%					
ADYNOVATE/ADYNOVI	15.3	14.2	14.3	14.3	15.4	0.6%	-3.3%	14.6	2.6%	-0.9%	-2.1%	15.9	11.5%	6.1%	0.6%					
FEIBA *3	12.9	7.7	13.7	10.3	11.4	-11.3%	-12.6%	8.8	13.7%	10.1%	-4.0%	8.8	-35.6%	-38.8%	-18.2%					
RECOMBINATE	3.7	3.2	3.5	2.9	3.7	-0.9%	-3.7%	2.6	-18.5%	-21.6%	-12.0%	3.3	-7.0%	-13.6%	-12.6%					
HEMOFIL/IMMUNATE/ IMMUNINE*3	4.4	4.9	3.9	5.4	3.3	-25.6%	-29.4%	5.1	3.0%	2.4%	-13.1%	5.2	33.5%	22.1%	-2.4%					
Other PDT Products *3 *4	0.9	0.8	0.9	0.9	0.9	-1.1%	-10.2%	1.1	31.3%	24.6%	6.4%	1.1	23.0%	15.8%	9.6%					
Others *5	5.9	5.4	5.8	6.0	6.9	16.4%	10.4%	6.6	22.5%	17.1%	13.6%	7.7	32.2%	24.7%	17.4%					
<b>Hereditary Angioedema</b>	<b>38.3</b>	<b>34.6</b>	<b>33.4</b>	<b>33.0</b>	<b>39.0</b>	<b>1.8%</b>	<b>-1.7%</b>	<b>35.2</b>	<b>1.8%</b>	<b>-2.2%</b>	<b>-1.9%</b>	<b>43.5</b>	<b>30.1%</b>	<b>21.1%</b>	<b>5.4%</b>					
TAKHZYRO	23.2	20.5	22.1	20.8	25.5	9.6%	6.0%	22.1	7.6%	3.2%	4.7%	30.9	39.5%	29.6%	13.2%					
FIRAZYR	8.1	7.1	5.0	6.7	6.9	-15.1%	-18.3%	7.5	5.9%	2.1%	-8.8%	7.1	43.9%	35.0%	2.2%					
CINRYZE *3	5.9	6.1	5.2	4.6	5.6	-5.7%	-9.2%	4.6	-24.3%	-27.3%	-18.4%	4.5	-14.2%	-19.8%	-18.8%					
KALBITOR	1.1	0.9	1.1	0.8	1.1	2.8%	0.8%	1.1	14.0%	9.9%	5.1%	1.0	-11.5%	-17.9%	-3.1%					
<b>Others</b>	<b>—</b>	<b>—</b>	<b>—</b>	<b>—</b>	<b>—</b>	<b>-</b>	<b>-</b>	<b>—</b>	<b>—</b>	<b>-</b>	<b>-</b>	<b>0.2</b>	<b>-</b>	<b>-</b>	<b>-</b>					

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 Generic name: pantoprazole

\*3 PDT products

\*4 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*5 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRI and Other Hemophilia.

[Table of Contents](#)

(Bn JPY)	FY20 Reported				FY21 Reported & Underlying Growth														
	Q1	Q2	Q3	Q4	YOY			YOY				YOY				YOY			
					Q1	Reported	Underlying	Q2	Reported	Underlying	YTD Underlying	Q3	Reported	Underlying	YTD Underlying	Q4	Reported	Underlying	YTD Underlying
<b>PDT Immunology</b>	<b>105.3</b>	<b>100.6</b>	<b>107.1</b>	<b>107.3</b>	<b>107.2</b>	<b>1.8%</b>	<b>-1.8%</b>	<b>130.8</b>	<b>30.0%</b>	<b>24.6%</b>	<b>11.1%</b>	<b>125.2</b>	<b>16.9%</b>	<b>8.7%</b>	<b>10.3%</b>				
immunoglobulin *1	85.1	77.6	85.4	86.8	81.6	-4.1%	-6.9%	99.7	28.6%	24.2%	8.0%	97.0	13.6%	6.1%	7.3%				
albumin *1	13.0	15.6	15.0	14.0	17.8	36.8%	26.4%	24.0	53.8%	42.2%	35.0%	19.7	31.4%	19.4%	29.7%				
Others *1 *4	7.2	7.5	6.7	6.5	7.8	9.1%	6.0%	7.1	-4.6%	-8.1%	-1.2%	8.5	25.7%	17.6%	4.8%				
<b>Oncology</b>	<b>108.0</b>	<b>102.1</b>	<b>108.4</b>	<b>98.0</b>	<b>121.4</b>	<b>12.4%</b>	<b>8.7%</b>	<b>112.3</b>	<b>10.0%</b>	<b>6.6%</b>	<b>7.7%</b>	<b>125.4</b>	<b>15.7%</b>	<b>9.2%</b>	<b>8.2%</b>				
VELCADE	24.2	25.8	25.9	25.2	30.1	24.6%	22.1%	25.0	-3.3%	-6.7%	7.1%	29.3	13.4%	5.3%	6.5%				
LEUPLIN/ENANTONE	27.4	22.5	25.4	20.1	26.2	-4.3%	-8.8%	27.6	23.0%	18.8%	3.6%	28.4	11.7%	7.7%	5.0%				
NINLARO	22.9	21.4	23.5	19.5	24.4	6.3%	2.0%	21.4	0.0%	-4.4%	-1.1%	24.9	6.1%	-1.1%	-1.1%				
ADCETRIS	15.1	15.5	13.8	15.0	17.2	14.2%	8.8%	16.9	9.3%	5.7%	7.2%	17.6	27.7%	22.2%	11.9%				
ICLUSIG	9.2	7.6	9.4	7.9	10.4	12.3%	10.0%	7.5	-1.6%	-5.0%	3.2%	8.8	-6.2%	-13.0%	-2.7%				
VECTIBIX	6.2	5.7	6.5	5.4	6.2	0.1%	0.1%	6.6	15.0%	15.0%	7.3%	6.6	1.7%	1.7%	5.3%				
ALUNBRIG	2.0	2.3	2.2	2.3	3.1	54.4%	47.3%	3.1	38.8%	33.0%	39.7%	3.9	75.6%	64.9%	48.4%				
Others	0.9	1.3	1.7	2.4	3.8	300.0%	307.2%	4.1	225.6%	242.2%	270.0%	5.8	234.8%	205.3%	241.7%				
<b>Neuroscience</b>	<b>106.9</b>	<b>100.9</b>	<b>107.3</b>	<b>102.2</b>	<b>113.4</b>	<b>6.1%</b>	<b>2.9%</b>	<b>120.3</b>	<b>19.2%</b>	<b>15.7%</b>	<b>9.1%</b>	<b>128.9</b>	<b>20.1%</b>	<b>11.7%</b>	<b>10.0%</b>				
VYVANSE/ELVANSE	66.0	66.6	69.8	69.1	79.2	20.0%	15.6%	80.1	20.2%	14.9%	15.2%	85.7	22.8%	13.5%	14.6%				
TRINTELLIX	16.9	18.1	17.7	16.2	17.9	5.9%	4.0%	22.2	22.7%	18.7%	11.6%	23.0	29.6%	21.0%	14.8%				
INTUNIV	5.6	3.3	5.9	5.6	3.3	-42.5%	-49.5%	4.2	26.6%	17.8%	-25.1%	5.0	-13.9%	-19.5%	-22.9%				
ADDERALL XR	5.3	3.7	4.4	4.4	3.9	-24.9%	-27.4%	5.7	52.9%	46.8%	3.5%	6.3	44.4%	33.5%	13.5%				
ROZEREM	3.0	2.9	3.6	2.5	3.2	6.9%	7.1%	3.1	5.7%	5.9%	6.5%	3.1	-12.6%	-12.4%	-0.6%				
Others *5	10.0	6.3	6.0	4.4	5.9	-41.2%	-41.1%	5.1	-19.3%	-3.5%	-27.7%	5.7	-4.0%	-8.0%	-21.8%				
<b>Others *2</b>	<b>139.8</b>	<b>152.0</b>	<b>153.5</b>	<b>128.7</b>	<b>241.6</b>	<b>72.8%</b>	<b>9.7%</b>	<b>118.2</b>	<b>-22.3%</b>	<b>10.0%</b>	<b>9.8%</b>	<b>122.3</b>	<b>-20.3%</b>	<b>12.0%</b>	<b>10.6%</b>				
AZILVA-F *3	20.9	19.1	22.9	19.4	22.6	8.6%	8.6%	17.7	-7.2%	-7.2%	1.1%	19.7	-13.8%	-13.8%	-4.4%				
LOTRIGA	8.1	7.6	8.8	7.3	7.8	-3.0%	-3.0%	8.2	8.5%	8.2%	2.4%	8.7	-1.3%	-1.6%	1.0%				
AIPHAGAN	4.0	3.7	4.6	3.7	4.6	15.0%	15.0%	3.8	3.3%	3.3%	9.4%	3.6	-20.7%	-20.7%	-1.9%				
FOSRENOL	3.2	3.3	3.7	3.3	3.4	4.7%	-3.2%	3.6	9.9%	4.5%	0.7%	3.2	-13.6%	-19.4%	-6.6%				
ACTOVEGIN	1.7	3.2	3.4	2.4	3.2	87.2%	83.3%	3.5	8.8%	3.8%	31.8%	4.3	29.4%	14.9%	24.7%				

\*1 PDT products

\*2 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21Q1 Reported.

\*3 The figures include the amounts of fixed dose combinations.

\*4 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*5 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL

Product Forecasts

(Bn JPY)	FY20 Reported Annual	Disclosed on May 11, 2021				Disclosed on October 28, 2021				Disclosed on February 3, 2022			
		FY21 Reported Forecasts			FY21 Underlying Growth Forecasts	FY21 Reported Forecasts			FY21 Underlying Growth Forecasts	FY21 Reported Forecasts			FY21 Underlying Growth Forecasts
		Annual	YOY			Annual	YOY			Annual	YOY		
<b>GI</b>	<b>777.8</b>	<b>878.0</b>	<b>100.2</b>	<b>13 %</b>	<b>10%</b>	<b>870.0</b>	<b>92.2</b>	<b>12 %</b>	<b>9%</b>	<b>883.0</b>	<b>105.2</b>	<b>14 %</b>	<b>8%</b>
ENTYVIO	429.3	538.0	108.7	25 %	22%	538.0	108.7	25 %	22%	542.0	112.7	26 %	19%
TAKECAB-F *1	84.8	94.0	9.2	11 %	11%	100.0	15.2	18 %	17%	101.0	16.2	19 %	18%
GATTEX/REVESTIVE	64.6	79.0	14.4	22 %	20%	74.0	9.4	15 %	12%	76.0	11.4	18 %	12%
DEXILANT	55.6	54.0	-1.6	-3 %	-6%	42.0	-13.6	-24 %	-27%	49.0	-6.6	-12 %	-17%
PANTOLOC/CONTROLOC*2	43.1	37.0	-6.1	-14 %	-19%	37.0	-6.1	-14 %	-19%	39.0	-4.1	-10 %	-14%
LIALDA/MEZAVANT	25.5	19.0	-6.5	-25 %	-25%	19.0	-6.5	-25 %	-25%	21.0	-4.5	-18 %	-20%
PENTASA	23.1	19.0	-4.1	-18 %	-20%	19.0	-4.1	-18 %	-20%	19.0	-4.1	-18 %	-20%
AMITIZA	21.2	5.0	-16.2	-76 %	-79%	5.0	-16.2	-76 %	-79%	4.0	-17.2	-81 %	-80%
RESOLOR/MOTTEGRITY	11.2	12.0	0.8	7 %	-1%	12.0	0.8	7 %	-1%	12.0	0.8	7 %	-1%
ALOFISEL	0.8	3.0	2.2	283 %	238%	3.0	2.2	283 %	238%	2.0	1.2	155 %	167%
Others	18.6	18.0	-0.6	-3 %	-8%	21.0	2.4	13 %	6%	18.0	-0.6	-3 %	-15%
<b>Rare Diseases</b>	<b>591.7</b>												
<b>Rare Metabolic</b>	<b>162.6</b>	<b>173.0</b>	<b>10.4</b>	<b>6 %</b>	<b>2%</b>	<b>173.0</b>	<b>10.4</b>	<b>6 %</b>	<b>2%</b>	<b>177.0</b>	<b>14.4</b>	<b>9 %</b>	<b>5%</b>
ELAPRASE	68.8	71.0	2.2	3 %	-1%	71.0	2.2	3 %	-1%	74.0	5.2	8 %	4%
REPLAGAL	51.8	56.0	4.2	8 %	3%	56.0	4.2	8 %	3%	56.0	4.2	8 %	5%
VPRIV	38.5	41.0	2.5	6 %	5%	41.0	2.5	6 %	5%	42.0	3.5	9 %	5%
NATPARANATPAR	3.6	5.0	1.4	41 %	38%	5.0	1.4	41 %	38%	5.0	1.4	41 %	35%
<b>Rare Hematology</b>	<b>289.8</b>	<b>273.0</b>	<b>-16.8</b>	<b>-6 %</b>	<b>-10%</b>	<b>273.0</b>	<b>-16.8</b>	<b>-6 %</b>	<b>-10%</b>	<b>279.0</b>	<b>-10.8</b>	<b>-4 %</b>	<b>-8%</b>
ADVATE	128.5	176.0	-10.6	-6 %	-10%	176.0	-10.6	-6 %	-10%	178.0	-8.6	-5 %	-9%
ADYNOVATE/ADYNOVI	58.1	35.0	-9.5	-21 %	-26%	35.0	-9.5	-21 %	-26%	38.0	-6.5	-15 %	-17%
FEIBA *3	44.5	12.0	-1.4	-10 %	-10%	12.0	-1.4	-10 %	-10%	12.0	-1.4	-10 %	-11%
RECOMBINATE	13.4	17.0	-1.7	-9 %	-13%	17.0	-1.7	-9 %	-13%	17.0	-1.7	-9 %	-10%
HEMOFIL/IMMUNATE/IMMUNINE*3	18.7	5.0	1.5	44 %	41%	5.0	1.5	44 %	41%	4.0	0.5	16 %	19%
Other PDT Products *3 *4	3.5	28.0	4.8	21 %	15%	28.0	4.8	21 %	15%	30.0	6.8	29 %	17%
Others *5	23.2												
<b>Hereditary Angioedema</b>	<b>139.3</b>	<b>0% to +10%</b>			<b>0% to +10%</b>	<b>0% to +10%</b>			<b>0% to +10%</b>	<b>+10% to +20%</b>			<b>0% to +10%</b>
TAKHZYRO	86.7	+20% to +30%			+20% to +30%	+20% to +30%			+20% to +30%	+20% to +30%			+10% to +20%
FIRAZYR	26.8	15.0	-11.8	-44 %	-46%	21.0	-5.8	-22 %	-24%	28.0	1.2	4 %	1%
CINRYZE *3	21.9	17.0	-4.9	-22 %	-23%	17.0	-4.9	-22 %	-23%	18.0	-3.9	-18 %	-22%
KALBITOR	3.9	2.0	-1.9	-49 %	-40%	2.0	-1.9	-49 %	-40%	4.0	0.1	2 %	-1%

\*1 The figures include the amounts of fixed dose combinations and blister packs.

\*2 Generic name: pantoprazole

\*3 PDT products

\*4 Other PDT products in Rare Hematology include BEBULIN, PROTHROMPLEX and Factor VII.

\*5 Other in Rare Hematology include VONVENDI, OBIZUR, RIXUBIS, AGRYLIN/XAGRID, OCTOFACTOR, COAGIL-VII, INNONAFACOR, and Other Hemophilia.

Average FX rates for FY2020: 1 USD = 106 JPY, 1 Euro = 123 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.6 JPY, 1 CNY = 15.5 JPY

Assumption of FX rates for FY2021 Reported Forecasts (Disclosed on May 11, 2021 and October 28, 2021): 1 USD = 108 JPY, 1 Euro = 131 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.9 JPY, 1 CNY = 16.8 JPY

Assumption of FX rates for FY2021 Reported Forecasts (Disclosed on February 3, 2022): 1 USD = 111 JPY, 1 Euro = 131 JPY, 1 RUB = 1.5 JPY, 1 BRL = 20.7 JPY, CNY = 17.3 JPY

Assumption of FX rates for FY2021 Underlying Forecasts: 1 USD = 106 JPY, 1 Euro = 123 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.6 JPY, 1 CNY = 15.5 JPY

(Bn JPY)	FY20 Reported	Disclosed on May 11, 2021				Disclosed on October 28, 2021				Disclosed on February 3, 2022			
		FY21 Reported Forecasts			FY21 Underlying Growth Forecasts	FY21 Reported Forecasts			FY21 Underlying Growth Forecasts	FY21 Reported Forecasts			FY21 Underlying Growth Forecasts
	Annual	YOY		Annual		YOY		Annual		YOY			
<b>PDT Immunology</b>	<b>420.4</b>	<b>+10% to +20%</b>			<b>+10% to +20%</b>	<b>+10% to +20%</b>			<b>+10% to +20%</b>	<b>+10% to +20%</b>			
immunoglobulin *1	334.9	+5% to +10%			+5% to +10%	+5% to +10%			+5% to +10%	+10% to +20%			
albumin *1	57.6	+>30%			+>30%	+>30%			+>30%	+>30%			
Others *1 *4	27.9	0% to +10%			0% to +10%	0% to +10%			0% to +10%	0% to +10%			
<b>Oncology</b>	<b>416.5</b>	<b>455.0</b>	<b>38.5</b>	<b>9%</b>	<b>7%</b>	<b>461.0</b>	<b>44.5</b>	<b>11%</b>	<b>8%</b>	<b>473.0</b>	<b>56.5</b>	<b>14%</b>	<b>9%</b>
VELCADE	101.1	83.0	-18.1	-18%	-20%	95.0	-6.1	-6%	-7%	108.0	6.9	7%	2%
LEUPLIN/ENANTONE	95.4	104.0	8.6	9%	7%	104.0	8.6	9%	7%	105.0	9.6	10%	7%
NINLARO	87.4	97.0	9.6	11%	8%	93.0	5.6	6%	4%	96.0	8.6	10%	4%
ADCETRIS	59.4	70.0	10.6	18%	14%	70.0	10.6	18%	14%	69.0	9.6	16%	12%
ICLUSIG	34.2	39.0	4.8	14%	11%	39.0	4.8	14%	11%	37.0	2.8	8%	3%
VECTIBIX	23.8	22.0	-1.8	-8%	-7%	22.0	-1.8	-8%	-7%	23.0	-0.8	-3%	-1%
ALUNBRIG	8.8	16.0	7.2	82%	80%	16.0	7.2	82%	80%	15.0	6.2	70%	59%
Others	6.4	24.0	17.6	276%	256%	22.0	15.6	245%	226%	20.0	13.6	213%	185%
<b>Neuroscience</b>	<b>417.3</b>	<b>434.0</b>	<b>16.7</b>	<b>4%</b>	<b>2%</b>	<b>450.0</b>	<b>32.7</b>	<b>8%</b>	<b>6%</b>	<b>474.0</b>	<b>56.7</b>	<b>14%</b>	<b>9%</b>
VYVANSE/ELVANSE	271.5	293.0	21.5	8%	5%	309.0	37.5	14%	11%	324.0	52.5	19%	13%
TRINTELLIX	68.9	82.0	13.1	19%	17%	82.0	13.1	19%	17%	82.0	13.1	19%	14%
INTUNIV	20.4	17.0	-3.4	-17%	-20%	17.0	-3.4	-17%	-20%	17.0	-3.4	-17%	-19%
ADDERALL XR	17.8	10.0	-7.8	-44%	-45%	13.0	-4.8	-27%	-28%	18.0	0.2	1%	-3%
ROZEREM	12.0	11.0	-1.0	-8%	-3%	11.0	-1.0	-8%	-3%	12.0	-0.0	-0%	2%
Others *5	26.7	21.0	-5.7	-21%	-17%	18.0	-8.7	-33%	-30%	21.0	-5.7	-21%	-17%
<b>Others *2</b>	<b>574.1</b>	<b>-10% to 0%</b>			<b>-10% to 0%</b>	<b>-10% to 0%</b>			<b>-10% to 0%</b>	<b>-10% to 0%</b>			<b>0% to +10%</b>
AZILVA-F *3	82.2	68.0	-14.2	-17%	-16%	68.0	-14.2	-17%	-16%	73.0	-9.2	-11%	-10%
LOTRIGA	31.8	29.0	-2.8	-9%	-8%	29.0	-2.8	-9%	-8%	31.0	-0.8	-2%	1%
AIPHAGAN	15.9	12.0	-3.9	-25%	-22%	12.0	-3.9	-25%	-22%	14.0	-1.9	-12%	-6%
FOSRENOL	13.5	11.0	-2.5	-18%	-17%	11.0	-2.5	-18%	-17%	12.0	-1.5	-11%	-16%
ACTOVEGIN	10.7	11.0	0.3	3%	7%	11.0	0.3	3%	7%	12.0	1.3	12%	10%

\*1 PDT products

\*2 The 133.0 billion JPY selling price of the sale of diabetes portfolio in Japan is included in FY21 Reported.

\*3 The figures include the amounts of fixed dose combinations.

\*4 Other in PDT Immunology include ARALAST, GLASSIA, CEPROTIN, ANTITHROMBIN III, KENKTSU-NONTHRON and others

\*5 Other in Neuroscience include REMINYL, COPAXONE, AZILECT, MYDAYIS, BUCCOLAM, EQUASYM and CARBATROL

Average FX rates for FY2020: 1 USD = 106 JPY, 1 Euro = 123 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.6 JPY, 1 CNY = 15.5 JPY

Assumption of FX rates for FY2021 Reported Forecasts (Disclosed on May 11, 2021 and October 28, 2021): 1 USD = 108 JPY, 1 Euro = 131 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.9 JPY, 1 CNY = 16.8 JPY

Assumption of FX rates for FY2021 Reported Forecasts (Disclosed on February 3, 2022): 1 USD = 111 JPY, 1 Euro = 131 JPY, 1 RUB = 1.5 JPY, 1 BRL = 20.7 JPY, CNY = 17.3 JPY

Assumption of FX rates for FY2021 Underlying Forecasts: 1 USD = 106 JPY, 1 Euro = 123 JPY, 1 RUB = 1.4 JPY, 1 BRL = 19.6 JPY, 1 CNY = 15.5 JPY

**Exchange Rate**

Average Exchange Rates vs. JPY				Impact of 1% depreciation of yen from January 2022 to March 2022 (Disclosed on February 3, 2022)			
CURRENCY	(yen)			(100 million yen)			
	FY20Q3 YTD (Apr-Dec)	FY21Q3 YTD (Apr-Dec)	FY21 Assumption (Apr-Mar) (Disclosed on February 3, 2022)	Revenue	Core Operating Profit	Operating Profit	Net Profit
USD	106	111	111	+30.1	+11.8	+5.2	+4.0
EUR	122	131	131	+8.0	-4.4	-6.2	-6.5
RUB	1.4	1.5	1.5	+1.1	+0.9	+0.8	+0.8
CNY	15.3	17.2	17.3	+2.2	+1.4	+1.4	+1.4
BRL	19.7	20.7	20.7	+1.0	+0.6	+0.5	+0.5

**CAPEX, depreciation and amortization and impairment losses**

<b>(Bn JPY)</b>	FY20	FY20Q3 YTD	FY21Q3 YTD	YOY		FY21 Forecasts (Disclosed on February 3, 2022)
Capital expenditures*	236.5	124.5	134.2	9.7	7.8%	210.0 -260.0
Tangible assets	111.2	75.0	87.7	12.6	16.8%	
Intangible assets	125.3	49.5	46.5	-2.9	-5.9%	
* Cash flow base						
Depreciation and amortization	558.0	419.3	428.4	9.0	2.2%	562.0
Depreciation of tangible assets* (A)	124.4	101.5	99.6	-2.0	-1.9%	
Amortization of intangible assets (B)	433.6	317.8	328.8	11.0	3.5%	
Of which Amortization associated with products (C)	405.3	304.6	309.1	4.5	1.5%	412.0
Of which Amortization excluding intangible assets associated with products (D)	28.3	13.2	19.7	6.5	49.0%	
* Excluding depreciation for investment assets.						
Depreciation and amortization (excluding intangible assets associated with products) (A)+(D)	152.7	114.8	119.3	4.5	4.0%	150.0
Impairment losses	25.5	10.1	14.7	4.5	44.9%	
Impairment losses associated with products	16.6	3.0	14.6	11.6	382.9%	40.0
Amortization and impairment losses on intangible assets associated with products	421.9	307.6	323.6	16.1	5.2%	452.0



### 3. Reconciliation

#### FY2021 Q3YTD Reconciliation from Reported Revenue to Core/Underlying Revenue

(Billion JPY)	Q3YTD		vs. PY	
	FY2020	FY2021		
<b>Reported Revenue</b>	<b>2,427.5</b>	<b>2,695.7</b>	<b>+268.2</b>	<b>+ 11.0%</b>
Sale of Japan diabetes portfolio*2	—	(133.0)	(133.0)	-5.5pp
<b>Core Revenue</b>	<b>2,427.5</b>	<b>2,562.7</b>	<b>+135.1</b>	<b>+ 5.6%</b>
FX effects*1				-4.8pp
Divestitures*2				+6.3pp
Regional portfolio				+4.6pp
Japan diabetes portfolio				+1.0pp
TACHOSIL				+0.5pp
Others				+0.1pp
<b>Underlying Revenue Growth</b>				<b>+ 7.1%</b>

\*1 FX adjustment applies plan rate to both periods.

\*2 Major adjustments are as follows:

- Revenue of select over-the-counter and non-core products in Asia Pacific is excluded from FY2020 Q3 YTD as the divestiture was completed in November 2020.
- Revenue of select non-core prescription pharmaceutical products predominantly in Europe is excluded from FY2020 Q3 YTD as the divestiture was completed in December 2020.
- Revenue of select over-the-counter and non-core products in Latin America is excluded from FY2020 Q3 YTD as the divestiture was completed in January 2021.
- Net sales from TACHOSIL, a surgical patch, are excluded from FY2020 Q3 YTD as the divestiture was completed in January 2021.
- Revenue of select over-the-counter and non-core products predominantly in Europe is excluded from FY2020 Q3 YTD as the divestiture was completed in March 2021.
- Revenue of the former subsidiary, Takeda Consumer Healthcare Company Limited is excluded from FY2020 Q3 YTD as the divestiture was completed in March 2021.
- Net sales from a portfolio of diabetes products in Japan (NESINA, LIOVEL, INISYNC and ZAFATEK) are excluded from FY2020 Q3 YTD as the divestiture was completed at the beginning of April 2021. In addition, the non-recurring item of the 133.0 billion JPY selling price as the result of the completion of the divestiture is excluded from FY2021 Q3 YTD.
- Revenue of select non-core prescription pharmaceutical products in China is excluded from both FY2021 Q3 YTD and FY2020 Q3 YTD as the divestiture was publicly announced and had been expected to complete within FY2021 H1. It is now expected to complete in FY2021 H2.

**FY2021 Q3YTD Reconciliation from Reported to Core/Underlying Core**

**FY2021 Q3YTD**

(Billion JPY)	REPORTED	REPORTED TO CORE ADJUSTMENTS							CORE	CORE TO UNDERLYING CORE ADJ.		UNDERLYING GROWTH	
		Amortization of intangible assets	Impairment of intangible assets	Other operating income/expenses	Sale of Japan diabetes portfolio	Irish Tax Assessment *1	TEVA JV related accounting adjustments	Others		FX	Divestitures		
Revenue	2,695.7				(133.0)				2,562.7	(112.1)	(14.0)	+7.1 %	
Cost of sales	(798.5)				0.6			33.1	(764.7)	35.4	4.1		
Gross Profit	1,897.3				(132.4)			33.1	1,798.0	(76.7)	(9.8)		
SG&A expenses	(662.9)				1.0			2.8	(659.1)	29.8	0.0		
R&D expenses	(382.5)							1.6	(380.9)	16.1	(0.0)		
Amortization of intangible assets	(309.1)	309.1							—				
Impairment losses on intangible assets	(14.6)		14.6						—				
Other operating income	34.3			(33.2)					—				
Other operating expenses	(100.0)			100.0					—				
Operating profit	462.5	309.1	14.6	66.9	(131.4)			(1.1)	37.5	757.9	(30.7)	(9.8)	+5.4 %
Margin	17.2 %									29.6 %			29.4 %*2
Financial income/expenses	(100.6)							11.6	(89.0)	8.3			
Equity income/loss	(5.3)							6.6	2.4	3.8	0.2		
Profit before tax	356.6	309.1	14.6	66.9	(131.4)			5.5	51.5	672.7	(22.2)	(9.8)	
Tax expenses	(115.1)	(68.9)	(3.6)	(17.5)	40.2	64.6	(1.7)	(49.1)	(151.1)	5.0	2.9		
Non-controlling interests	(0.1)								(0.1)	(0.0)	0.0		
Net profit	241.4	240.2	10.9	49.4	(91.2)	64.6	3.8	2.3	521.5	(17.2)	(6.9)		
EPS (yen)	154								333	(10)	(4)		+9.9 %
Number of shares (millions)	1,567								1,567				1,563

\*1 A tax charge of 64.6 billion JPY for tax and interest, net of 0.5 billion JPY of associated tax benefit, arising from tax assessment involving Irish taxation of the break fee Shire received from AbbVie in connection with the terminated offer to acquire Shire made by AbbVie in 2014.

\*2 Underlying Core Operating Profit Margin.

**FY2020 Q3YTD Reconciliation from Reported to Core/ Underlying Core**
**FY2020 Q3YTD**

(Billion JPY)	REPORTED	REPORTED TO CORE ADJUSTMENTS					CORE	CORE TO UNDERLYING CORE ADJ.		UNDERLYING GROWTH
		Amortization of intangible assets	Impairment of intangible assets	Other operating income/expenses	TEVA JV related accounting adjustments	Others		FX	Divestitures	
Revenue	2,427.5						2,427.5	3.2	(155.1)	+1.1 %
Cost of sales	(740.9)					73.8	(667.0)	(7.2)	43.9	
Gross Profit	1,686.7					73.8	1,760.5	(4.0)	(111.3)	
SG&A expenses	(641.3)			0.0		(0.3)	(641.5)	1.5	12.4	
R&D expenses	(342.5)			(0.4)		4.5	(338.4)	0.9	0.6	
Amortization of intangible assets	(304.6)	304.6					—			
Impairment losses on intangible assets	(3.0)		3.0				—			
Other operating income	118.5			(57.3)	(1.1)	(60.2)	—			
Other operating expenses	(155.1)			136.4		18.7	—			
Operating profit	358.7	304.6	3.0	78.9	(1.1)	36.6	780.6	(1.6)	(98.3)	+8.5 %
Margin	14.8 %						32.2 %			29.9 %*
Financial income/expenses	(115.4)					17.2	(98.2)	6.1	(0.0)	
Equity income/loss	(8.0)					16.2	3.0	(0.0)	(0.0)	
Profit before tax	235.4	304.6	3.0	78.9	15.1	48.6	685.5	4.5	(98.3)	
Tax expenses	(56.3)	(68.5)	(0.6)	(14.1)	(4.6)	(21.4)	(165.5)	(1.1)	27.4	
Non-controlling interests	(0.1)						(0.1)	0.0	0.0	
Net profit	178.9	236.1	2.5	64.8	10.5	27.1	519.8	3.4	(70.9)	
EPS (yen)	115						333	3	(45)	+4.5 %
Number of shares (millions)	1,562						1,562			1,558

\* Underlying Core Operating Profit Margin.

## Free Cash Flow

(BN JPY)	FY2020 Q3 YTD	FY2021 Q3 YTD	vs. PY	
<b>Net profit</b>	<b>179.0</b>	<b>241.5</b>	<b>62.5</b>	<b>+34.9 %</b>
Depreciation, amortization and impairment loss	430.4	445.5	15.1	
Decrease (increase) in trade working capital	(48.9)	41.2	90.1	
Income taxes paid	(174.7)	(107.2)	67.5	
Tax refunds and interest on tax refunds received	28.4	6.1	(22.2)	
Other	195.8	120.3	(75.5)	
<b>Net cash from operating activities</b>	<b>610.0</b>	<b>747.5</b>	<b>137.6</b>	<b>+22.6 %</b>
Adjustment for cash temporarily held by Takeda on behalf of third parties <sup>1</sup>	—	47.0	47.0	
Acquisition of PP&E	(75.0)	(87.7)	(12.6)	
Proceeds from sales of PP&E	42.8	0.4	(42.4)	
Acquisition of intangible assets	(49.5)	(46.5)	2.9	
Acquisition of investments	(9.5)	(7.6)	1.9	
Proceeds from sales and redemption of investments	73.7	16.1	(57.7)	
Proceeds from sales of business, net of cash and cash equivalents divested	125.0	2.1	(122.8)	
<b>Free Cash Flow</b>	<b>717.5</b>	<b>671.3</b>	<b>(46.2)</b>	<b>(6.4)%</b>

1. Adjustment refers to cash temporarily held by Takeda on behalf of third parties related to vaccine operations and the trade receivables sales program.

**FY2021 Q3 YTD NET PROFIT TO ADJUSTED EBITDA BRIDGE**

(BN JPY)	FY2020 Full Year (Apr-Mar)	FY2020 Q3 YTD (Apr - Dec)	FY2021 Q3 YTD (Apr - Dec)	FY2021 Q3 YTD LTM* <sup>1</sup> (Jan-Dec)
<b>Net profit</b>	<b>376.2</b>	<b>179.0</b>	<b>241.5</b>	<b>438.7</b>
Income tax expenses	(9.9)	56.3	115.1	48.8
Depreciation and amortization	559.7	420.3	430.9	570.3
Interest expense, net	129.0	99.7	86.7	116.0
<b>EBITDA</b>	<b>1,054.9</b>	<b>755.3</b>	<b>874.2</b>	<b>1,173.8</b>
Impairment losses	25.5	10.1	14.7	30.0
Other operating expense (income), net, excluding depreciation and amortization and other miscellaneous expenses (non-cash item)	(74.5)	26.4	59.5	(41.3)
Finance expense (income), net, excluding interest income and expense, net	14.1	15.7	13.9	12.3
Share of loss on investments accounted for under the equity method	(0.1)	8.0	5.3	(2.8)
Other adjustments:	131.4	102.3	(46.6)	(17.6)
Non-core expense related to COVID-19	14.0	8.8	7.2	12.4
Sale of Japan diabetes portfolio	—	—	(131.4)	(131.4)
Impact on profit related to fair value step up of inventory in Shire acquisition	79.4	68.0	24.8	36.1
Acquisition costs related to Shire	1.9	0.0	—	1.9
Other costs* <sup>2</sup>	36.1	25.5	52.9	63.5
<b>Adjusted EBITDA</b>	<b>1,151.3</b>	<b>917.9</b>	<b>920.9</b>	<b>1,154.4</b>
EBITDA from divested products* <sup>3</sup>				(10.3)
<b>Adjusted EBITDA (LTM)</b>				<b>1,144.1</b>

\*<sup>1</sup> LTM represents Last Twelve Months (January 2021 - December 2021). Calculated by subtracting FY2020 Q3 YTD from FY2020 Full Year and adding FY2021 Q3 YTD.

\*<sup>2</sup> Includes adjustments for non-cash equity-based compensation expense and other one time non-cash expense.

\*<sup>3</sup> Represents adjustments for EBITDA from divested products which are removed as part of LTM Adjusted EBITDA.

## Net Debt to Adjusted EBITDA

### FY2021 Q3 YTD

#### NET DEBT/ADJUSTED EBITDA RATIO

(BN JPY)	FY2021 Q3 YTD
Cash and cash equivalents*1	595.9
Book value debt on the balance sheet	(4,354.9)
Hybrid bond 50% equity credit	250.0
FX adjustment*2	117.2
Gross debt*3	(3,987.7)
<b>Net cash (debt)</b>	<b>(3,391.9)</b>
<b>Net debt/Adjusted EBITDA ratio</b>	<b>3.0 x</b>
<b>Adjusted EBITDA</b>	<b>1,144.1</b>

#### NET INCREASE (DECREASE) IN CASH

(BN JPY)	FY2020 Q3 YTD	FY2021 Q3 YTD	vs. PY	
Net cash from operating activities	610.0	747.5	137.6	+22.6 %
Acquisition of PP&E	(75.0)	(87.7)		
Proceeds from sales of PP&E	42.8	0.4		
Acquisition of intangible assets	(49.5)	(46.5)		
Acquisition of investments	(9.5)	(7.6)		
Proceeds from sales and redemption of investments	73.7	16.1		
Acquisition of business, net of cash and cash equivalents acquired	—	(49.7)		
Proceeds from sales of business, net of cash and cash equivalents divested	125.0	2.1		
Net increase (decrease) in short-term loans and commercial papers	(85.0)	(0.0)		
Repayment of long-term loans	(792.5)	(414.1)		
Proceeds from issuance of bonds	1,179.5	249.3		
Repayment of bonds	(596.6)	(220.9)		
Interest paid	(84.2)	(84.9)		
Dividends paid	(274.7)	(273.0)		
Others	(72.1)	(82.4)		
Net increase (decrease) in cash	(8.1)	(251.4)	(243.3)	(2,999.7)%

\*1 Includes short-term investments which mature or become due within one year from the reporting date and excludes cash temporarily held by Takeda on behalf of third parties related to vaccine operations and the trade receivables sales program.

\*2 FX adjustment refers to change from month-end rate to average rate used for non-JPY debt calculation outstanding at the beginning of the period to match with adjusted EBITDA (which is calculated based on average rates). New non-JPY debt incurred and existing non-JPY debt redeemed during the reporting period are translated to JPY at relevant spot rates as of the relevant date.

\*3 Bonds and loans of current and non-current liabilities. 250Bn yen reduction in debt due to 500Bn yen hybrid bond issuance in June 2019, given that the hybrid bond qualifies for 50% equity credit for leverage purposes. Includes cash and non cash adjustments to debt book-value. Non cash adjustments include changes dues to debt amortization and FX impact.

**FY2020 Q4 (Full Year)**

<b>NET DEBT/ADJUSTED EBITDA RATIO</b>	
(BN JPY)	FY2020
Cash and cash equivalents*1	790.7
Book value debt on the balance sheet	(4,635.4)
Hybrid bond 50% equity credit	250.0
FX adjustment*2	165.2
Gross debt*3	(4,220.2)
<b>Net cash (debt)</b>	<b>(3,429.4)</b>
<b>Net debt/Adjusted EBITDA ratio</b>	<b>3.2 x</b>
<b>Adjusted EBITDA</b>	<b>1,083.5</b>

<b>NET INCREASE (DECREASE) IN CASH</b>			
(BN JPY)	FY2019	FY2020	vs. PY
Net cash from operating activities	669.8	1,010.9	+341.2 +50.9%
Acquisition of PP&E	(127.1)	(111.2)	
Proceeds from sales of PP&E	12.6	46.5	
Acquisition of intangible assets	(90.6)	(125.3)	
Acquisition of investments	(7.6)	(12.6)	
Proceeds from sales and redemption of investments	49.4	74.6	
Acquisition of business, net of cash and cash equivalents acquired	(4.9)	—	
Proceeds from sales of business, net of cash and cash equivalents divested	461.5	530.4	
Net increase (decrease) in short-term loans and commercial papers	(351.2)	(149.0)	
Repayment of long-term loans	(137.4)	(792.5)	
Proceeds from issuance of bonds	496.2	1,179.5	
Repayment of bonds	(563.6)	(859.2)	
Interest paid	(127.2)	(107.3)	
Dividends paid	(282.6)	(283.4)	
Others	(40.6)	(85.3)	
Net increase (decrease) in cash	(43.3)	316.1	+359.4 —

\*1 Includes short-term investments which mature or become due within one year from the reporting date and excludes deposits restricted to certain vaccines operations.

\*2 FX adjustment refers to change from month-end rate to average rate used for non-JPY debt calculation, to match with adjusted EBITDA calculation.

\*3 Bonds and loans of current and non-current liabilities. 250Bn yen reduction in debt due to 500Bn yen hybrid bond issuance in June 2019, given that the hybrid bond qualifies for 50% equity credit for leverage purposes. Includes cash and non cash adjustments to debt book-value. Non cash adjustments include changes dues to debt amortization and FX impact.

**Reconciliation from Reported Operating Profit to Core Operating Profit - FY2021 Forecast**

(BN JPY)	REPORTED	REPORTED TO CORE ADJUSTMENTS					CORE
		Amortization of intangible assets	Impairment of intangible assets	Other operating income/expenses	Sale of Japan diabetes portfolio	Others	
<b>Revenue</b>	<b>3,510.0</b>				<b>(133.0)</b>		<b>3,377.0</b>
Cost of sales					0.6	36.2	
<b>Gross Profit</b>					<b>(132.4)</b>	<b>36.2</b>	
SG&A and R&D expenses					1.0	(3.8)	
Amortization of intangible assets	(412.0)	412.0					—
Impairment losses on intangible assets	(40.0)		40.0				—
Other operating income	48.0			(48.0)			—
Other operating expenses	(150.0)			150.0			—
<b>Operating profit</b>	<b>515.0</b>	<b>412.0</b>	<b>40.0</b>	<b>102.0</b>	<b>(131.4)</b>	<b>32.4</b>	<b>970.0</b>



### **Important Notice**

For the purposes of this notice, "report" 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 release. This report (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 report. 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 report 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 report, "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.

The product names appearing in this document are trademarks or registered trademarks owned by Takeda, or their respective owners.

### **Forward-Looking Statements**

This report and any materials distributed in connection with this report 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. These forward-looking statements are based on assumptions about many important factors, including the following, which could cause actual results to differ materially from those expressed or implied by the forward-looking statements: 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, including global health care reforms; challenges inherent in new product development, including uncertainty of clinical success and decisions of regulatory authorities and the timing thereof; uncertainty of commercial success for new and existing products; manufacturing difficulties or delays; fluctuations in interest and currency exchange rates; claims or concerns regarding the safety or efficacy of marketed products or product candidates; the impact of health crises, like the novel coronavirus pandemic, on Takeda and its customers and suppliers, including foreign governments in countries in which Takeda operates, or on other facets of its business; the timing and impact of post-merger integration efforts with acquired companies; the ability to divest assets that are not core to Takeda's operations and the timing of any such divestment(s); and other factors identified 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/sec-filings/> or at [www.sec.gov](http://www.sec.gov). Takeda does not undertake to update any of the forward-looking statements contained in this report 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 or statements of Takeda in this report may not be indicative of, and are not an estimate, forecast, guarantee or projection of Takeda's future results.

### **Certain Non-IFRS Financial Measures**

This report includes certain non-IFRS financial measures and targets. Takeda's management evaluates results and makes operating and investment decisions using both IFRS and non-IFRS measures included in this report. Non-IFRS results exclude certain income and cost items which are included in IFRS results. By including these non-IFRS measures, management intends to provide investors with additional information to further analyze Takeda's performance, core results and underlying trends. Non-IFRS results are not prepared in accordance with IFRS and non-IFRS information should be considered a supplement to, and not a substitute for, financial statements prepared in accordance with IFRS. Investors are encouraged to review the reconciliations of non-IFRS financial measures to their most directly comparable IFRS measures, which are on "*Supplementary Information - 3. Reconciliation.*"

### **Medical information**

This report 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").