# NXera ~

# Corporate Presentation March 2025 | Nxera Pharma Co., Ltd. (TSE: 4565)

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Agenda

- Business Overview
- Strategic Roadmap
- Our Pipeline
- Japan/APAC Business
- <sup>o5</sup> Our NxWave<sup>TM</sup> Platform
- Financial Results
- Appendix



# **Business** Overview

## Leading the Next Era of Medicine. From Japan, for Japan, and the world

World-leading NxWave<sup>™</sup> platform (UK), coupled with Japan's most effective development and commercial organization

# **Our Mission**

To accelerate the development of life-changing medicines, by investing in science and technology.

# **Our Vision**

To lead the next era of medicine.

From Japan, for Japan, and the world.

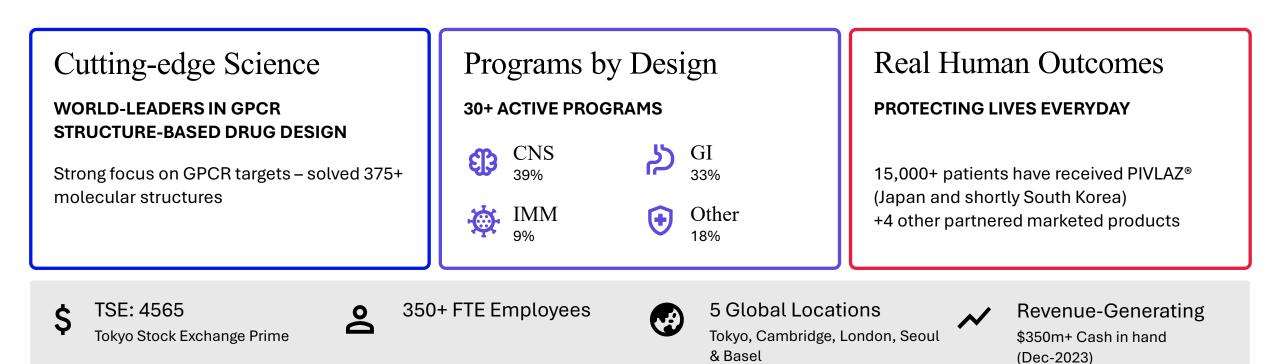
# Our Values

- Patients come first
- Innovation and teamwork
- Focus
- Speed and agility
- Operational excellence



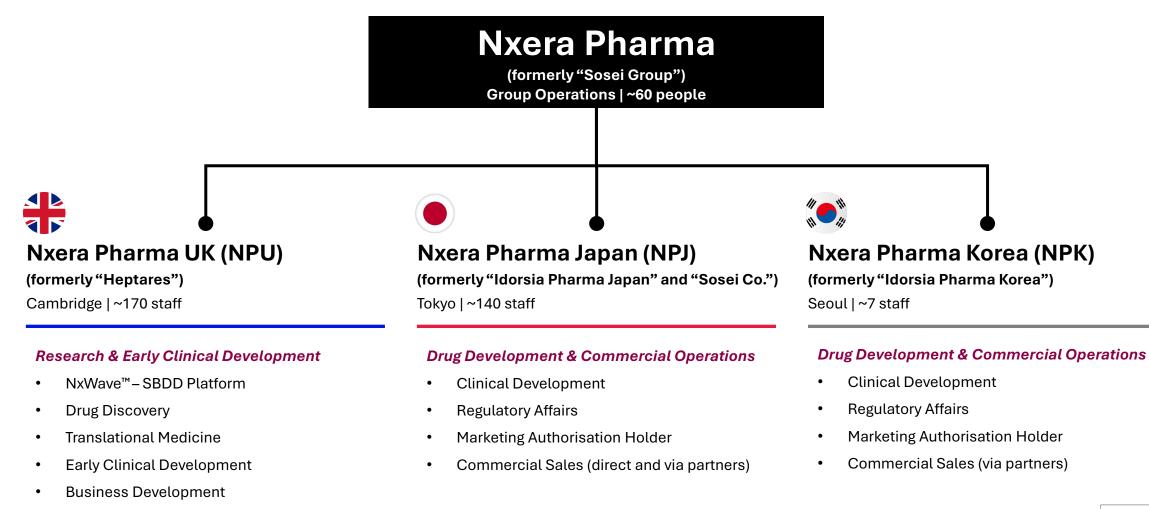
### We are Nxera Pharma

A technology-powered biopharma in pursuit of new specialty medicines to improve the lives of patients



### **Global Corporate Structure**

Over 350 team members employed across Japan, South Korea, UK and Switzerland



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JP/APAC

# Agile and decisive leadership team



# Strategic Roadmap

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# Our History

Strategic steps taken to build Nxera over the last two decades

2000s	2015	2023	2024
Launched a public company dedicated to <b>bringing innovation to</b> Japan ✓ IPO on TSE (MOTHERS) in 2004	<ul> <li>Out-licensed several programs to global pharma to generate profit, a cash reserve and a larger market valuation</li> <li>✓ 15+ partnered programs that generate upfront and milestone revenue (plus future royalties)</li> </ul>	<ul> <li>Elevated our status in the Tokyo</li> <li>Stock Exchange, improving access to institutional investors</li> <li>✓ Promotion to TSE (PRIME) segment in 2023</li> <li>✓ First public healthcare investment by the Japan Investment Corporation in 2023</li> </ul>	<b>EXERCISE</b> Launched new corporate branding: <b>Nxera Pharma Co</b>
<ul> <li>Made strategic acquisitions to bring steady revenue through groundbreaking medicines</li> <li>✓ \$186m acquisition of Arakis Limited in 2005</li> <li>✓ Royalty revenues from Breezhaler® medicines from 2012 to present</li> </ul>	<ul> <li>Invested in research-focused companies that could generate a continuous pipeline of new medicines</li> <li>✓ \$400m acquisition of Heptares Therapeutics Limited in 2015</li> </ul>	<ul> <li>Acquired a commercial-stage pharmaceutical company which provided an integrated platform for even greater sustainable revenue growth</li> <li>✓ \$466m acquisition of Idorsia Pharmaceuticals Japan and Korea</li> <li>✓ Rapidly growing revenues from sales of PIVLAZ<sup>®</sup></li> </ul>	With a vision to lead the next era of medicine. From Japan, for Japan, and the world.
ARAKIS	HEPTARES therapeutics	<b>IDOISIA</b> JAPAN KOREA	



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# To make our mission happen...

Accelerate the development of life-changing medicines

123Acquire or in-license<br/>multiple de-risked<br/>medicines for JapanInvest in our<br/>NxWave<sup>™</sup> platform<br/>to seed programsBuild a first-class<br/>technology<br/>environment

Focusing on these three areas is how we plan to make our mission happen as fast as possible



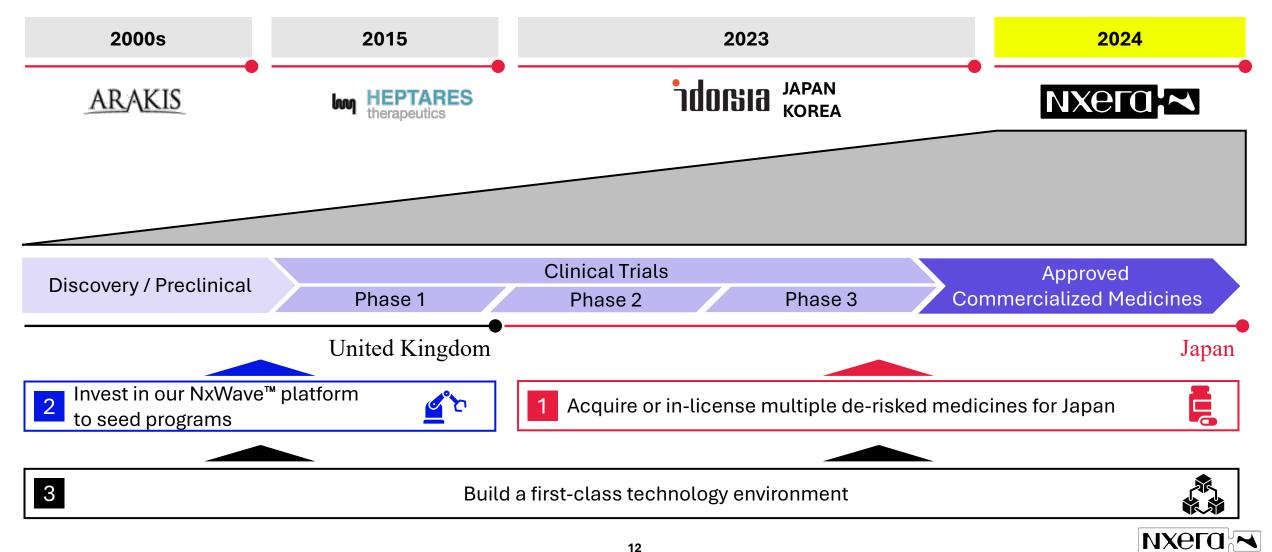
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# ...building a fully integrated biopharma from Japan

Accelerating growth to achieve our mission by leveraging business platform in Japan and UK



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# Priority objectives for FY2025

JPY 17 billion+ Net product sales (PIVLAZ<sup>®</sup> plus QUVIVIQ<sup>®</sup>)

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Acquire/in-license at least one late-stage medicine for Japan/APAC (ex-China)

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Execute <u>at least one</u> new major partnership, and initiate <u>at least one</u> new in-house Ph.2 study



Investment in systems and applications for efficiency and scalability

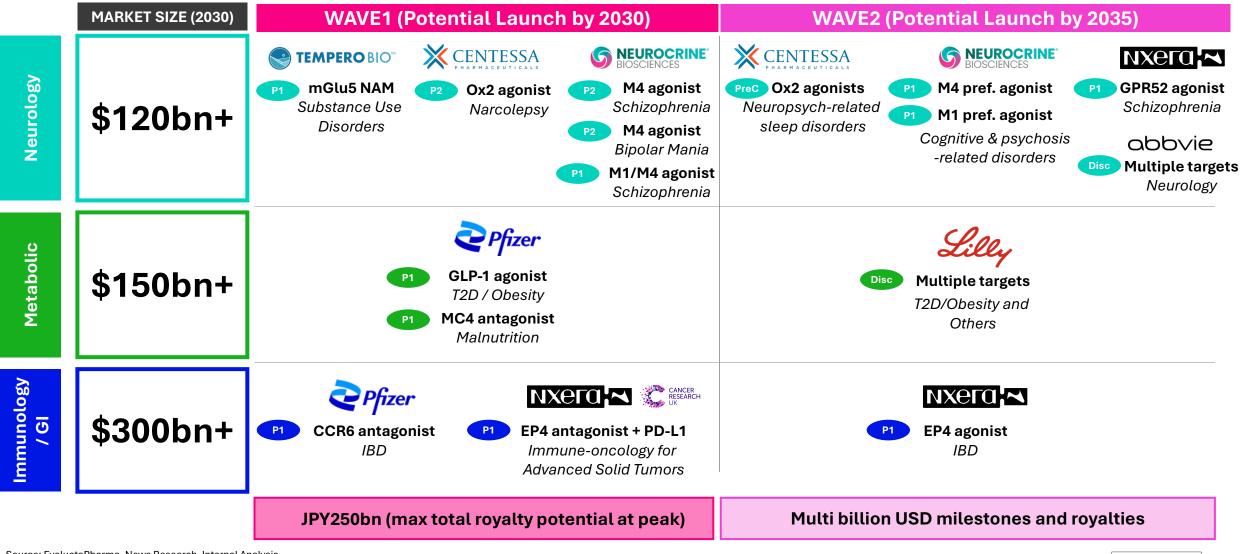


Positive operating profit under IFRS (if GPR52 option is exercised)



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Our Wave 1 and Wave 2 programs are positioned across fast growing areas of healthcare



Source: EvaluatePharma, News Research, Internal Analysis

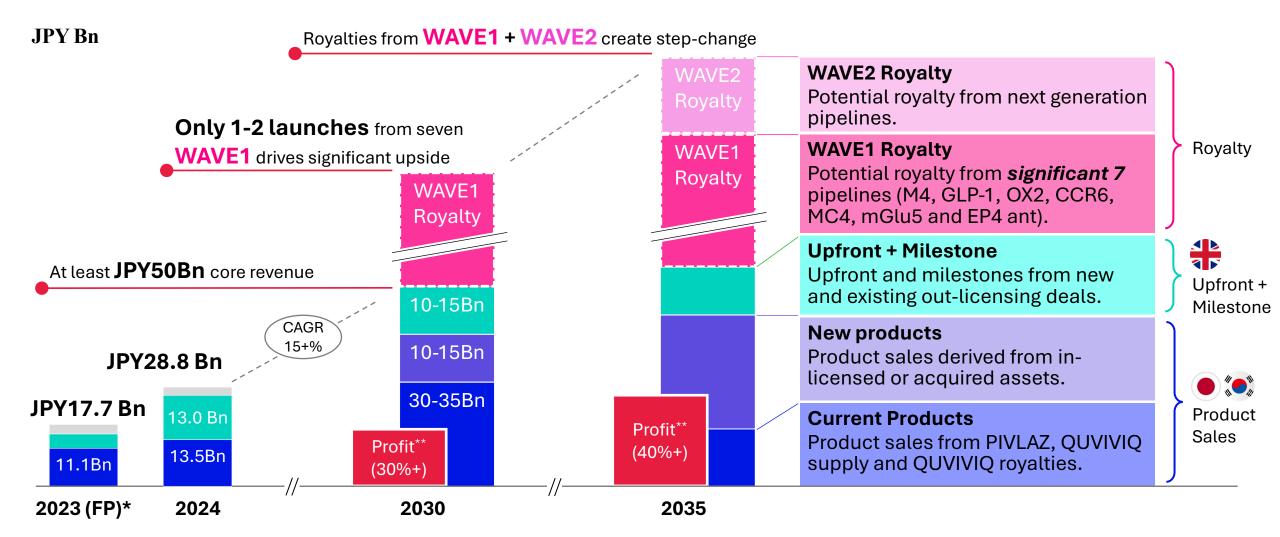
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Our 2030 vision is to build a high growth, highly profitable Japanese biopharma



Note: \* Revenue values are proforma the acquisition of Idorsia Pharmaceuticals Japan and Korea and reflect annual product sales of PIVLAZ in 2023.

\*\* WAVE1 and WAVE2 royalty is not included.



Our Pipeline Programs by Design

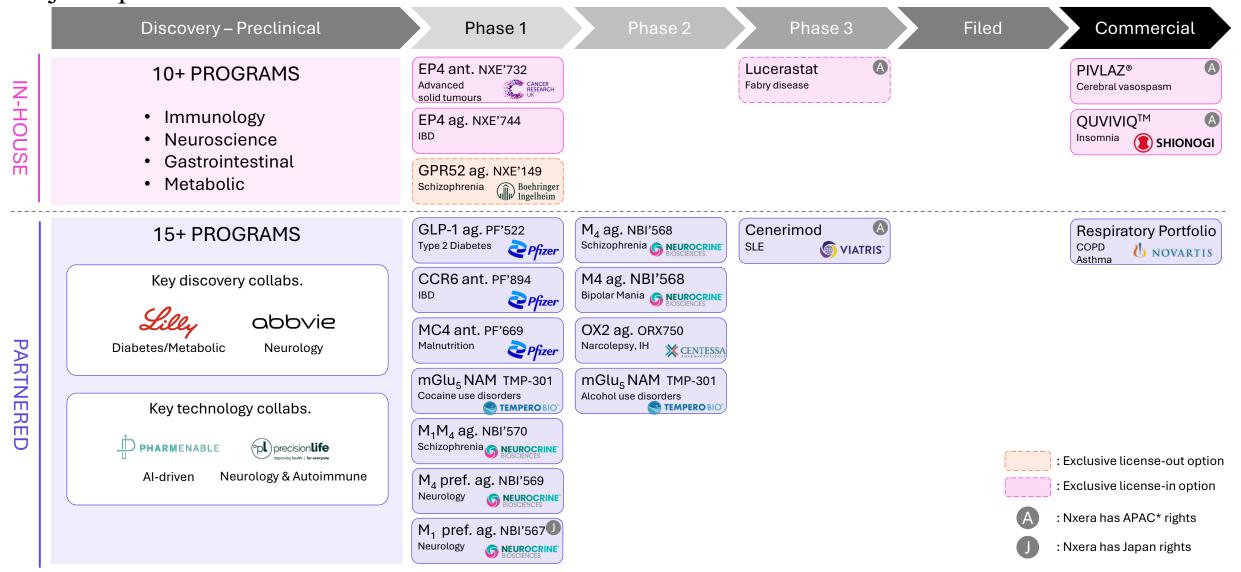


OADMAP PIPELINE JP/APAC

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# Major Pipeline Overview



Note: Pref. ag. : Preferring agonist

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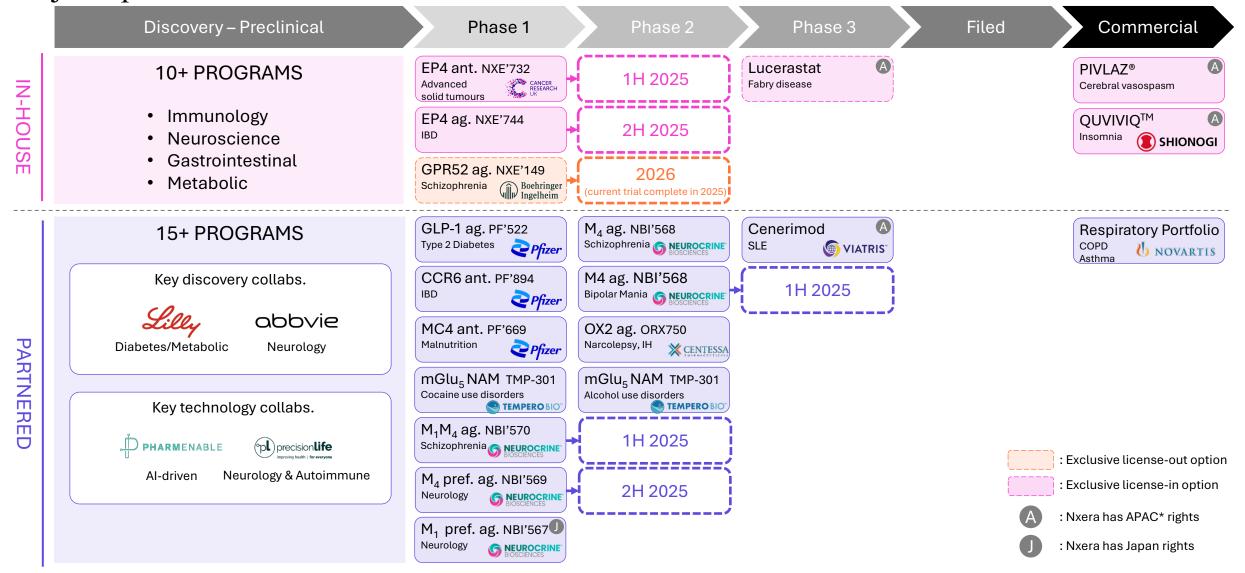
\*APAC (ex-China) territory includes Japan, South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

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# Major Pipeline Overview



Note: Pref. ag. : Preferring agonist

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\*APAC (ex-China) territory includes Japan, South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

# Looking ahead to potential catalysts in 2025<sup>\*</sup>

PROGRAM	PARTNER	TIMING	EVENT
TMP-301 (mGlu5 NAM)		H1 2025	Phase 2 study start in alcohol use disorder
Cenerimod (S1P1) / Lucerastat	ndorsia	H1 2025	Exclusive opt-in decision
NXE'732 (EP4 antagonist)		H1 2025	Phase 2a study start in Advancing Solid Tumors
NBI'568 (M4 agonist)	<b>SEUROCRINE</b> BIOSCIENCES	H1 2025	Phase 3 study start in Schizophrenia
NBI'568 (M4 agonist)		H2 2025	Phase 2 study start in Bipolar Mania
NBI'570 (M1/M4 agonist)	<b>BIOSCIENCES</b>	H2 2025	Phase 2 study start in Schizophrenia
NXE'744 (EP4 agonist)	NXELCI 🛪	H2 2025	Phase 2 study start in IBD
NXE'149 (GPR52 ag)	NXCIO Sochringer Ingelheim	H2 2025	Phase 1b completion
NXE'732 (EP4 antagonist)		H2 2025	Phase 1b topline data
ORX750 (OX2 agonist)		H2 2025	Phase 2a data across NT1, NT2, and IH
Multiple discovery collaboration progress	abbvie <i>Lilly</i>	2025	Progression through discovery stage
NBI'567 (M1 ago) / NBI'569 (M4 ago) / NBI'570 (M1/M4 ago)	<b>BIOSCIENCES</b>	2025	Phase 1 data readout
New global out-licenses		Anytime	Out licensing and/or discovery collabs
New Japan / APAC in-licenses		Anytime	Acquire/in-license late-stage medicines
QUVIVIQ™		Anytime	APAC out-licensing deals

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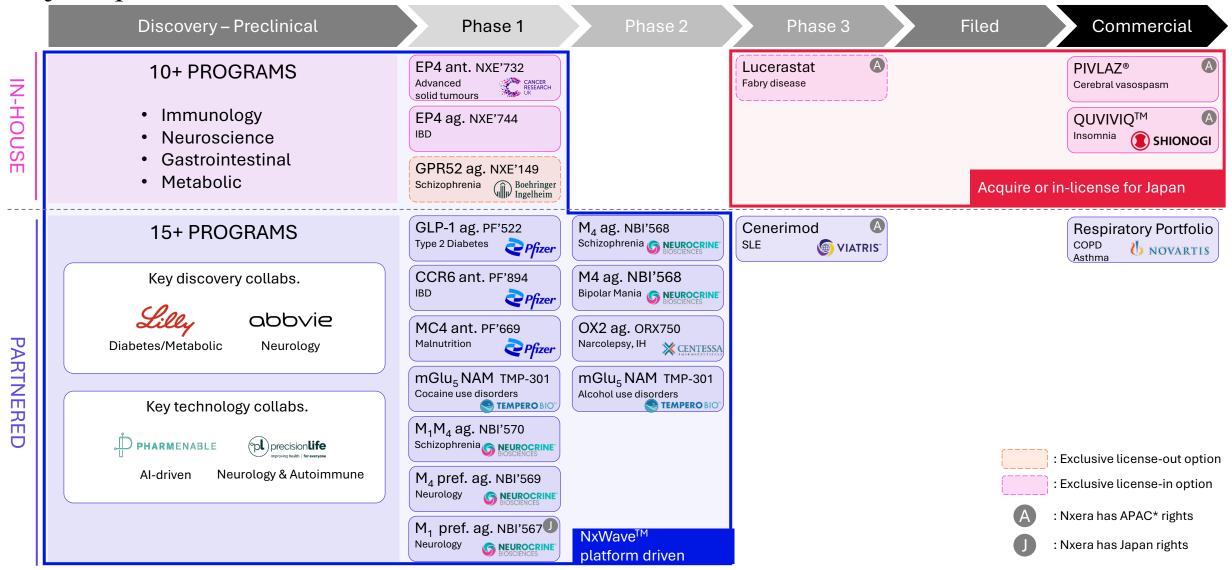
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### Major Pipeline Overview



Note: Pref. ag. : Preferring agonist

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\*APAC (ex-China) territory includes Japan, South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

# Key strategy for each business category

Maximize the value of each business and demonstrate synergies by conducting integrated development in future

	NxWave <sup>™</sup> platform driven <b></b> ℃	Acquire or in-license for Japan
Organic Growth	<ul> <li>Collaborate with existing partner to help them to progress pipeline licensed from us</li> <li>Execute at least one new high value collaboration and/or co-investment per year</li> </ul>	Maximize and optimize sales and profit for two major products (PIVLAZ <sup>®</sup> /QUVIVIQ <sup>™</sup> )
Strategic Growth	Collaborate/invest in new technologies with synergies	In-license late-stage products for clinical development and commercialization in Japan and APAC

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**NEUROCRINE**®

**X** CENTESSA

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# Delivering science with commercial potential

#### **Nxera's Commercialized Products**

1

Neurological disorders – diseases of ageing

### **PIVLAZ®**



 prevention of cerebral vasospasm in patients with Aneurysmal Subarachnoid Hemorrhage (aSAH) Partnered Products (Discovered by Nxera/with NxWave<sup>™</sup> tech)



Neurological disorders – psychiatric / cognition

– Muscarinic agonists

– Orexin 2 agonists

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Neurological disorders – QOL diseases - sleep

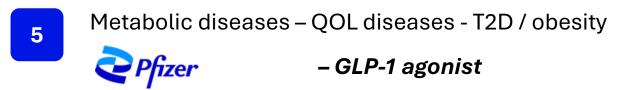
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Neurological disorders – quality of life diseases

# QUVIVIQ™



- treatment of adult patients with insomnia



# **JPY30–35bn** product sales by 2030 (plus, multiple other programs in discovery/development)

#### Up to JPY250bn royalty revenues at peak

(plus, multiple other programs in discovery/development)



# Japan/APAC Business

Deliver innovation to patients in Japan/APAC

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## Japan will serve as our base to expand across APAC markets

Japan is an attractive, established market with strong volumes

APAC is the second Tailwinds from near-Japan is the second High quality clinical largest pharma and regulatory highest growth term regulatory market (ex-China) pharma market changes environment Market size (USD bn) Market growth (CAGR %) 66 (2021)(2019 - 2027) Japan Phase 1 Drug Excellent access to 580 10.0% 8.5% Doctors/HCPs who evaluate **Clinical Trials No** 200 169 7.0% 8.0% novel drugs 6.0% Longer Needed for 5.5% 150 6.0% 4.0% **Global Clinical Trials** Typically achieve 85 4.0% 100 65 42 strong patient uptake 2.0% 50 " 0.0% Reduces drug loss and drug 0 APAC Latin Africa & Europe North US China Japan Germany France MHLW lag for Japan patients Middle America America East

Source: IQVIA Market Prognosis, Sep 2022; IQVIA Institute, Nov 2022.

APAC (ex-China) territory includes South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

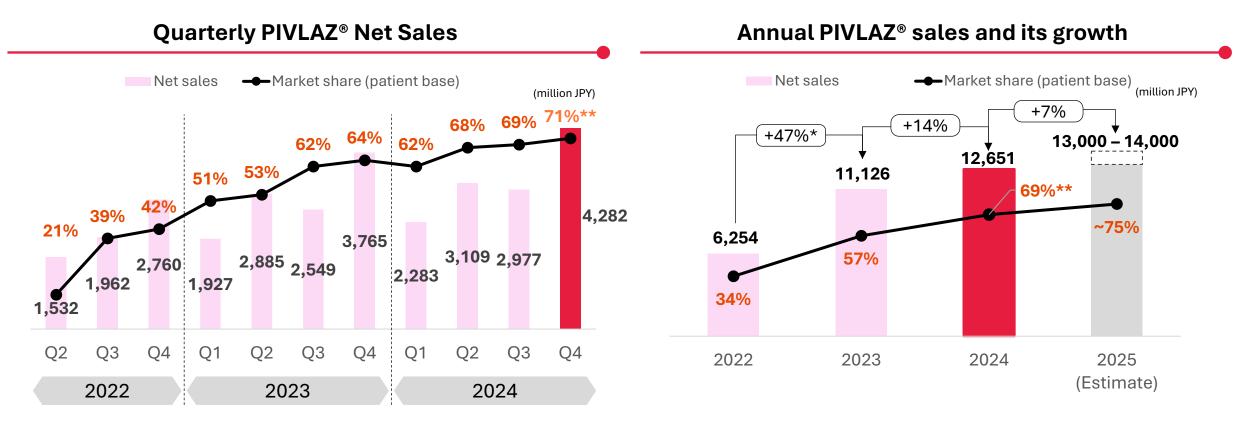


エンドセリン受容体拮抗薬

ビヴラッツ®点滴静注液

# PIVLAZ<sup>®</sup> (clazosentan, an endothelin A antagonist)

Our first commercially available product for the prevention of cerebral vasospasm in patients with Aneurysmal Subarachnoid Haemorrhage (aSAH)



PIVLAZ<sup>®</sup> has rapidly built awareness and is becoming the standard of care with neurosurgeons



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薬価基準収載

150mg

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FINANCIAL | OPERATIONAL HIGHLIGHTS

HLIGHTS JAPAN/APAC

R&D PROGRESS FY2025 OBJECTIVES

**クービビック®錠 25mg** 50mg

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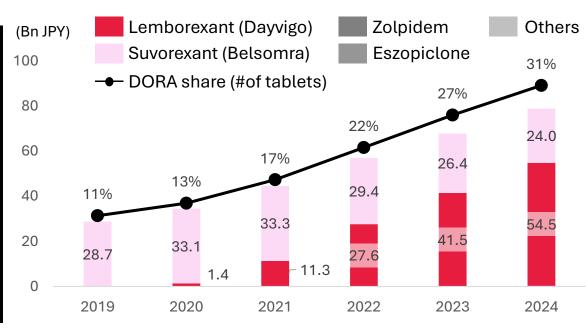
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薬価基準未収載

# QUVIVIQ<sup>TM</sup> (daridorexant, dual orexin antagonist "DORA")

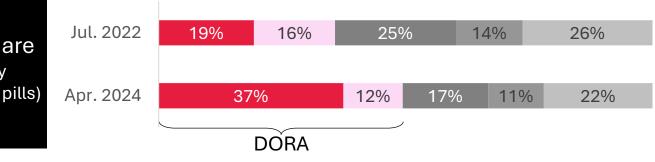
DORA is rapidly establishing its position in the treatment paradigm for insomnia

Sales and market share (NHI-base)



- DORAs are rapidly penetrating the insomnia treatment market in Japan, where traditional anti-anxiety and z-class drugs are not preferred by physicians
- Japan is one of the largest
   DORA markets globally –
   estimated at up to US\$1bn
- Together with partner Shionogi, we aim to provide a best-inclass product

#### Prescription share (Most frequently prescribed sleeping pills)

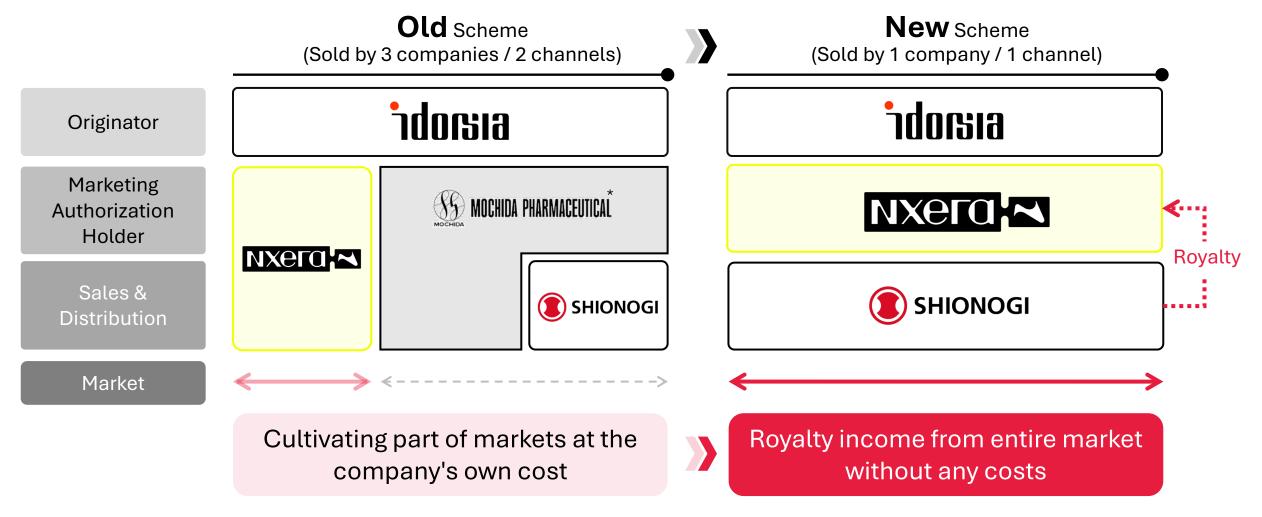


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**クービビック**®錠 25mg

# QUVIVIQ<sup>TM</sup> Business scheme change

SHIONOGI to Exclusively Handle Distribution and Sales Activities in Japan





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#### クービビック<sup>®</sup>錠 25mg In-house pipeline: QUVIVIQ<sup>TM</sup> JNDA approval received in Sep. 2024 and launched in Dec. 2024. Aim to be the best-in-class drug Unmet needs in insomnia About QUVIVIQ<sup>™</sup> **Dual Orexin** Alleviates excessive wakefulness through **Receptor Antagonist** strong inhibition of orexin receptors Nocturnal awakenings Recommended in the 2023 European Insomnia **European Guideline** Guidelines as the only orexin receptor antagonist that can be used <sup>1</sup> Rapid sleep onset T<sub>max</sub>: about 0.5-1.4 hour T<sub>1/2</sub>: about 6-9 hour Carry-over effects to the PK profile Significant improvement in next-day sleepiness next day after medication and daytime functioning confirmed in global phase 3 trials <sup>2</sup>

Aim to be the Best-in-class drug in DORA class

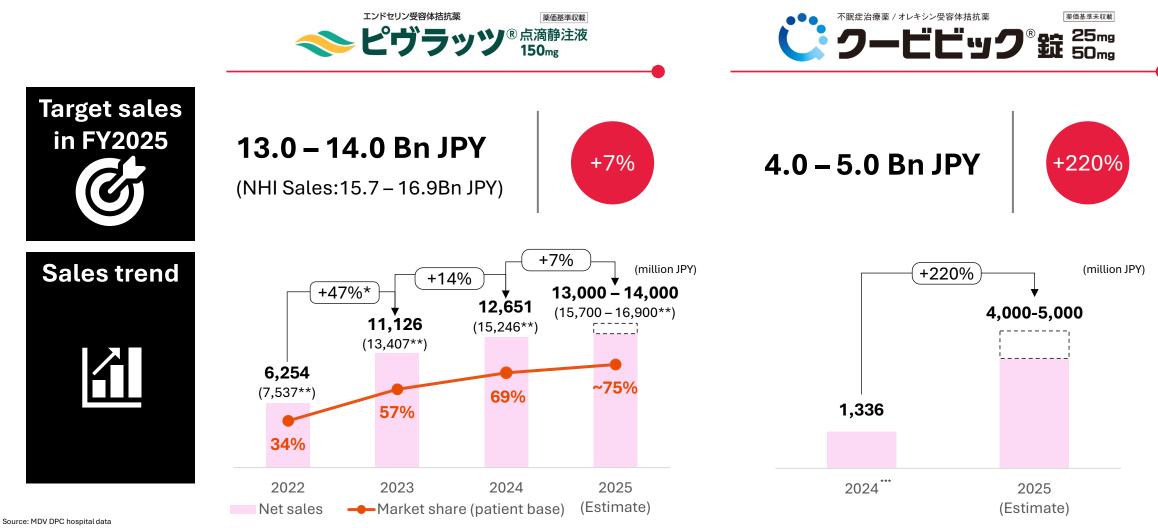


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## Full year product sales guidance

Target 13.0 - 14.0 Bn JPY (PIVLAZ) as net sales, and 4.0 - 5.0 Bn JPY (QUVIVIQ) as royalty and supply







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\*: Comparison of 2-4Q of 2022 and 2023, \*\* NHI sales, \*\*\* 2024 sales includes upfront, milestone, royalty and product supply while 2025 sales includes royalty and product supply

# Our NxWave<sup>TM</sup> Platform

Cutting-edge Science

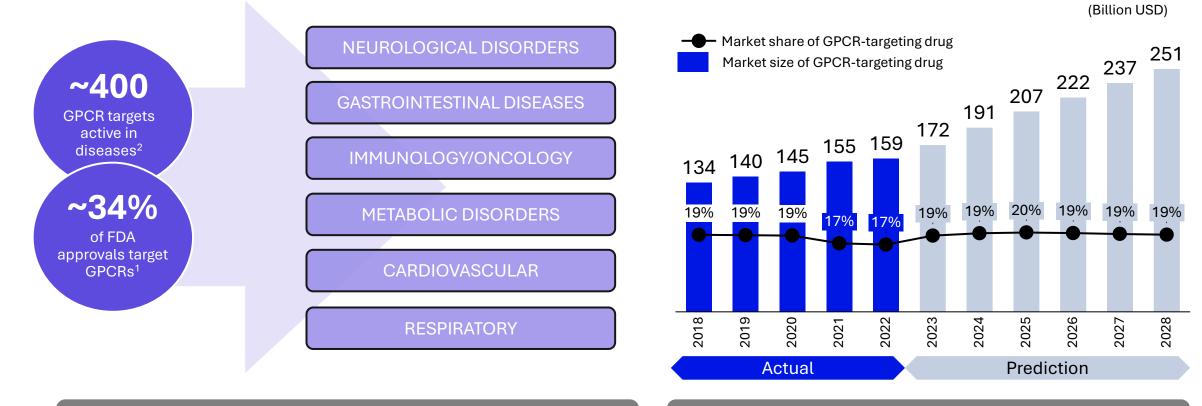
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# NxWave<sup>TM</sup> platform is focussed on drugging GPCRs

GPCRs are the largest family of drug discovery targets – comprising 1/3 of all FDA approved drugs



# GPCRs are active in a wide range of disease areas, and offer broad therapeutic potential

# Drugs that target GPCRs account for 20% of the entire pharmaceutical market

Soruce: <sup>1</sup> "Unexplored opportunities in the druggable human genome", Nature Reviews, 2016; <sup>2</sup> "Trends in GPCR in Drug Discovery – new agents, targets and indications", Nature Reviews, 2017, GPCRs as targets for approved drugs: How many targets and how many drugs? (2018), Evaluate Pharma, The IUPHAR/BPS Guide to PHARMACOLOGY



# GPCR: Large unmet needs and FIC opportunities

>650 First-in-class opportunities in GPCR-targeting drug

### Best-in-class opportunities (~120): Drugs are available



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Total ~800 drug opportunities (~400 GPCRs are thought to be drug targets)



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# NxWave<sup>TM</sup> platform enables faster, cheaper and more precise drug discovery

World-leading science and platform enables efficient drug discovery against difficult targets

	<b>Conventional drug discovery</b>	Our drug discovery	
Approach	Empirical design	Rational design (computer-based)	
Method	High Throughput Screening (HTS <sup>1</sup> )	Proprietary NxWave <sup>™</sup> Platform	
Period <sup>2</sup>	4.5 years on average	3.0 years on average	
Costs <sup>2</sup>	\$15 million	\$5 million	
Features <sup>3</sup>	Difficult to design drugs precisely – high development attrition rate	Execute more precise drug design – lower development attrition rate	
Target <sup>3</sup>	Difficult for GPCRs with unstable structures	Best for GPCRs with unstable structures	

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<sup>1</sup> HTS/High Throughput Screening is a method to find drug candidates by reacting tens of thousands to millions of compounds with drug targets using large machines and human hands.

<sup>2</sup> The period from target selection to preclinical testing. For conventional drug discovery, figures are taken from NATURE REVIEWS Drug Discovery (MARCH 2010).

<sup>3</sup> Precise drug design make clear the binding site of target, make easier to improve compound, create backups and redo – potentially increase the success rate. GPCR is most popular drug target which account for 30% of current drug target.





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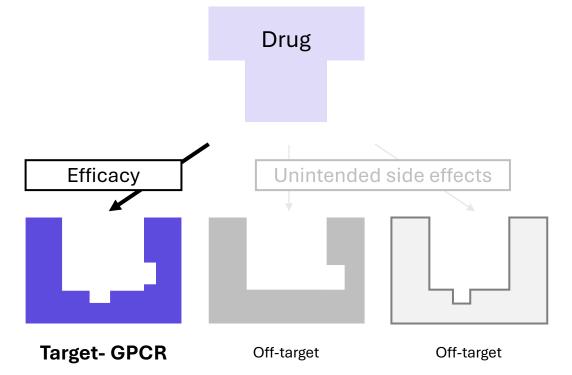


# Our platform enables precise design of GPCR models

Only by performing detailed structural analysis can we design great drugs.

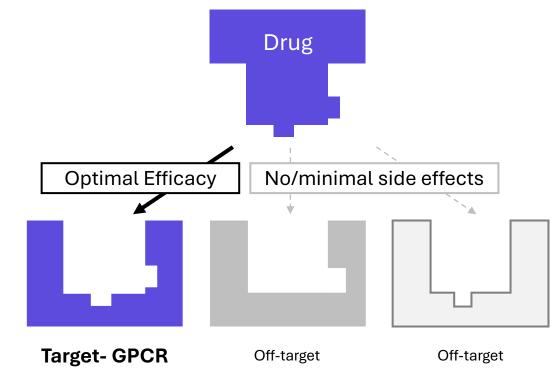
### Imprecise GPCR model: Standard Medicine

Poorly understood GPCRs (locks) led to suboptimal drugs (keys) being designed



### Precise GPCR model: Optimized Medicine

High selectivity enables to **optimize efficacy and minimize side effects** 





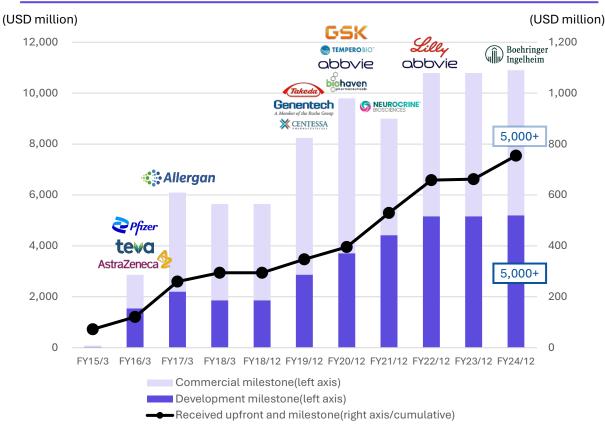
# Our track record of major licensing transactions speaks for itself...

Income from licensing provides a great source of non-dilutive financing to support our growth

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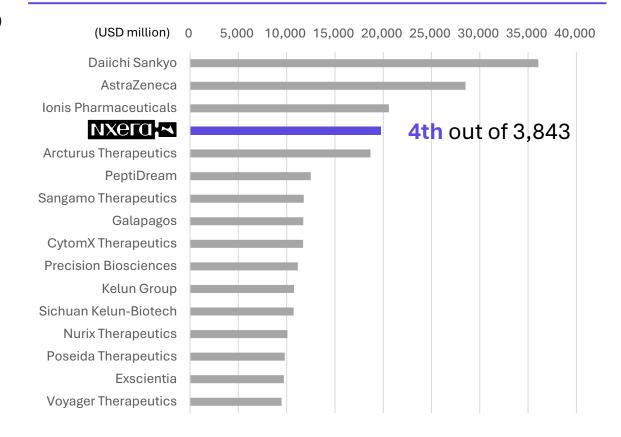
# Balance of potential milestone income from existing license agreements<sup>1</sup>



#### Top 15 pharmaceutical/biotech companies by license value<sup>2</sup> (cumulative total since 2015)

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<sup>1</sup> Balance as of the end of the fiscal year of only those currently under contract. TEVA and AbbVie (formerly Allergan), for which compounds were returned, are excluded from the balances from FY2018 and FY2021, respectively.
 <sup>2</sup> The figures are based on 'Licensing' category on third party's (EvaluatePharma's) proprietary database and therefore do not completely match the amounts shown in the LHS chart.
 Source: Company's data (LHS) and EvaluatePharma (as of 2024/10/17) (RHS)





Platform

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Platform

# ... hundreds of millions of dollars received, billions of dollars in potential to come

New collaboration and exclusive option to license agreement executed with Boehringer Ingelheim

Partner	Execution	Program	Therapeutic Area(s)	Upfront and Initial Milestones	Potential Total Milestone <sup>1</sup>
Boehringer Ingelheim	March 2024	Collaboration and exclusive option-to- license agreement for GPR52 agonist	Schizophrenia	€25m	€670m
Lilly	December 2022	Multi-target Collaboration	Diabetes and Metabolic	\$37m	\$800m
abbvie	August 2022	Multi-target Collaboration	Neurological disorders	\$80m	\$1.2bn
<b>DEUROCRINE</b> BIOSCIENCES	December 2021	Collaboration and license agreement for $M_4$ , $M_1$ and $M_1/M_4$ dual agonist	Neurological disorders	\$100m	\$2.6bn
GSK	December 2020	Collaboration and license agreement for GPR 35	Gastrointestinal, immunology	\$44m	\$480m
behaven	December 2020	Collaboration and license agreement for CGRP portfolio	Neurology	\$10m	\$380m
abbvie	June 2020	Discovery Collaboration and Option to License <sup>2</sup>	Inflammatory and Autoimmune	\$32m	\$400m
Takeda	August 2019	Multi-target Collaboration	Multiple; Initial focus on Gastrointestinal	\$26m	\$1.2bn
<b>Genentech</b> A Member of the Roche Group	July 2019	Multi-target Collaboration	Multiple	\$26m	\$1.0bn
<b>P</b> fizer	November 2015	Multi-target Collaboration	Multiple	-	\$1.8bn

<sup>1</sup>Potential option fees, development, regulatory and commercial milestone payments agreed at the time of transaction. Nxera is also eligible to receive tiered royalties ranging from high single digit to mid-teen percentage on future net sales of any products developed under the partnership.<sup>2</sup> AbbVie has the option to expand the collaboration by an additional three targets



#### M4 ago. (NBI'568) demonstrated competitive positive phase 2 data



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Once-daily 20 mg dose showed efficacy, and good safety / tolerability profile for schizophrenia patients.

	PANSS total score change	-18.2		
Clinically meaningful and	PANSS total score change vs. Placebo	-7.5 (p = 0.011)	Met primary and	
statistically significant efficacy (Once-daily 20 mg dose)	Effect size	0.61	additional endpoints and	
	Marder Factor score change vs Placebo:		demonstrated <u>efficacy</u> on both positive and	
	Positive	-3.0 (p=0.004)	negative symptoms	
	Negative	-1.9 (p=0.028)		
Generally safe and well-tolerated	Treatment discontinuation rate due to adverse events across all NBI'568 arms	5.0% (placebo: 4.3%)	NBI'568 showed <u>safety</u>	
across all doses tested	<ul> <li>GI and CV adverse event frequency (Cobenfy (BMS/Karuna): 3-5x (GI), ~4x (CV) vs. placebo)</li> </ul>	Similar to placebo	and tolerability for all doses	
	Received successful milestone of Ph2 trial	US\$ 35 m		
Rapidly advancing to	Ph3 clinical trial	begin in 1H 2025	Expanding potential of	
Phase 3 development	Additional Ph2 trial in Bipolar Mania	begin in 2H 2025	muscarinic agonist portfolio	
	Advancing follow-on compounds in muscarinic			

OVERVIEW

STRATEGIC ROADMAP

Source: Presentation of Neurocrine Sciences (Aug.28 2024), KarXT for Schizophrenia draft evidence report (Nov. 28, 2023)

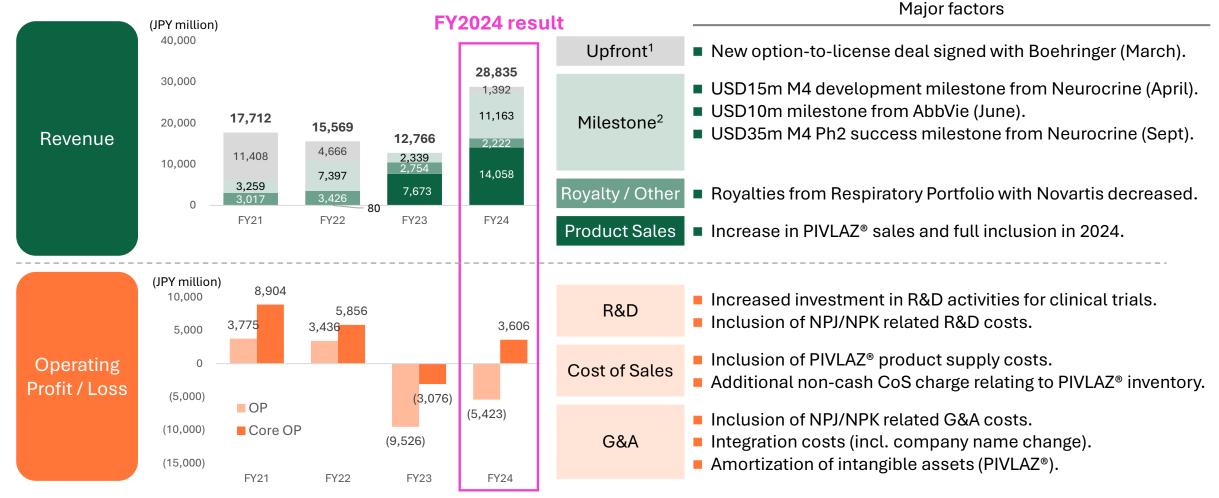


## **Financial Results**



#### Key financial indicators

NPJ/NPK product sales and cost base fully included in FY2024. Milestones increased due to progress of partnered programs



<sup>1</sup> Upfront fee revenue recognised at deal inception

<sup>2</sup> Milestone revenue recognised at milestone event + deferred revenue releases



#### Breakdown of 2024 results

Impact of Non-cash/Non-recurring costs was more significant in 2024 due to the inclusion of the Idorsia businesses

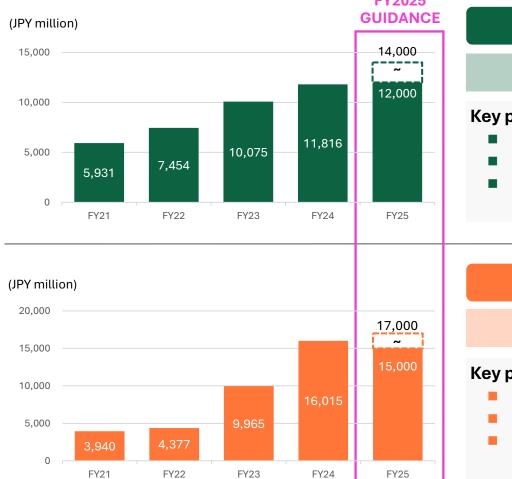
(JPY million)	NPC / NPU <sup>*1</sup>	NPJ / NPK <sup>*2</sup>	Consolidated P&L (Core)	Non-cash costs 🕀	Non-recurring Costs	Consolidated P&L (IFRS)			
Revenue	14,847	13,988	28,835		,	28,835			
Cost of Sales + SG&A	(7,015)	(8,963)	(15,978)	A PIVLA B (1,362) Amortization - Product IP	(2,401) AZ® inventory adjustment (1,160) Integration (2,730) Other	(23,630)			
R&D	(9,258)	(1,242)	(10,500)		(1,316)	(11,816)			
Other income	1,272	(23)	1,249	D	(60)	1,189			
OP/Core OP	(154)	3,760	Core OP 3,606	Total : 9,0	OP (5,423)				
M&A related costsAdditional CoS charge for PIVLAZ® stock which completed by 3Q 2024.B o Amortization of intangible assets (currently relates to PIVLAZ®). Annual charge to increase to c. JPY 1,800m per year from 2025.C o o oIntegration costs including IT system integration and Corporate rebranding. Will significantly decrease in 2025.									
Other	Amortization of other in	ntangible assets (e.g. II	P), depreciation (e.g. labo	ratory equipment), share-based	d payments and other re	estructuring costs.			

\*1 = Nxera Pharma Co. Ltd. (formerly Sosei Group Corporation) + Nxera Pharma UK Ltd (formerly Heptares Therapeutics Ltd.) + Sosei K.K
 \*2 = Nxera Pharma Japan (formerly Idorsia Pharmaceuticals Japan) + Nxera Pharma Korea (formerly Idorsia Pharmaceuticals Korea)



#### Full year cost guidance for FY2025

Slight increase in R&D expenses with progression of programs into later stages of development and in-licensing of one or more late-stage candidates. Lower to flat SG&A expenses through streamlining costs



FY2025	skpenses unough streamming costs
GUIDANCE	R&D expenses (IFRS basis)
14,000 ~ 12,000	JPY12,000 to JPY14,000m
FY25	<ul> <li>Key points in FY2025</li> <li>Incremental investment in platform technology.</li> <li>In-house programs (EP4 ant., EP4 ago., GPR52 ago.) moving into Ph1b - Ph2.</li> <li>Clinical development of one or more in-licensed late-stage assets in Japan.</li> </ul>
	S&M + G&A expenses (IFRS basis)
17,000	JPY15,000 to JPY17,000m
15,000	<ul> <li>Key points in FY2025</li> <li>Investment in technology to increase efficiency and deliver future growth.</li> <li>Increase in amortization as QUVIVIQ<sup>™</sup> has launched.</li> <li>Lower or flat SG&amp;A expenses vs. FY2024 through cost savings.</li> </ul>



APPENDIX



## Exclusive Opt-in Rights And ROFN/ROFR<sup>1</sup>

Option to develop up to five clinical programs for Japan and APAC (ex-China) from Idorsia

	Program	Mechanism of Action	Indication	Stage	Region
Exclusive Opt-in Right	Lucerastat	Glucosylceramide synthase inhibitor	Fabry disease	Phase 3	
	ACT-1004-1239	ACKR3 / CXCR7 antagonist	Multiple sclerosis and other demyelinating diseases	Phase 2*	
ROFR	ACT-1014-6470	C5aR1 antagonist	Immune-mediated disorders	Phase 1*	APAC (ex-China) <sup>2</sup>
/ROFN <sup>1</sup>	IDOR-1117-2520	Undisclosed	Immune-mediated disorders	Phase 1*	
	ACT-777991	CXCR3 antagonist	Recent-onset Type 1 diabetes	Phase 1*	

<sup>1</sup> ROFN/ROFR - Right of first negotiation / Right of first refusal

<sup>2</sup> Territories include Japan, South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

\* Global Phase

APPENDIX

Core Operating Profit/Loss – a financial indicator closer to the reality of our business

#### **Operating Profit**

"Core"

- Core Operating Profit/ Loss is a key financial indicator that highlights the underlying recurring cash generating capability of our business.
- Core Operating Profit/Loss is defined as IFRS Operating Profit + material Non-cash costs + material non-recurring costs
- Material Non-cash Costs include depreciation, amortization, share based payments and impairment.
- Material Non-recurring Costs include restructuring costs, M&A related professional fees and other material one-off items.

#### + Material Non-cash Costs

STRATEGIC ROADMAP

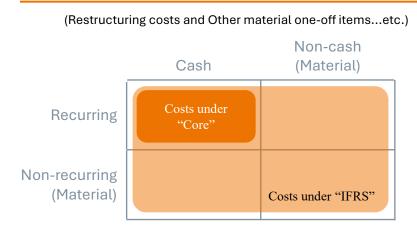
PIPELINE

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(Depreciation, Amortization, Share based payments, Impairment...etc.)

#### + Material Non-recurring Costs



## Operating Profit **"IFRS"**

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 Financial results recorded and prepared in accordance with International Financial Reporting Standards (IFRS)



#### Estimation of potential market size

Multi-billion USD annual peak sales potential for our post-pre-clinical pipeline

Cotogony	Indication <sup>2</sup>	Number of Patients –	Pe	eak Sales	Candidates	
Category	mulcation-		Market Size	Individual Products	Candidates	
	Dementia	~55 million	\$7.3 billion (2010)	\$3.9 billion (2009/Aricept)	M1 ag, M1/M4 ag	
Neuroscience	Schizophrenia	~20 million	\$20.7 billion (2011)	\$5.7 billion (2013/Abilify)	M4 ag, M1/M4 ag, GPR52 ag	
Neuroscience	Substance use disorders	~10.4 million <sup>1</sup>			mGlu5 NAM	
	Narcolepsy	~3 million	\$2.5 billion (2024)	\$1.4 billion (2024/Xywav)	OX2 ag	
	Cancer	~42 million	\$210.5 billion (2024)	\$28.7 billion (2024/Keytruda)	EP4 ant	
Immunology	IBD	~10 million	\$23.8 billion (2024)	\$6.2 billion (2022/Humira)	CCR6 ant, GPR35 ag, EP4 ag	
	Systemic Lupus Erythematosus	~5 million	\$2.7 billion (2024)	\$1.9 billion (2024/Benlysta)	Cenerimod	
Metabolism	T2DM/Obesity	~420 million	\$76.8 billion (2024)	\$18.2 billion (2024/Ozempic)	GLP1 ag	
	Anorexia	~10 million			MC4 ant	
	Total		~\$344 billion/year	~\$66 billion/year		

Source (Number of patients): World Health Organization, Evaluate Pharma, The European Federation of Crohn's & Ulcerative Colitis Associations (EFCCA), Narcolepsy Network, Inc., The Lupus Foundation of America, GBD 2015 Disease and Injury Incidence and Prevalence Collaborators (October 2016). "Global, regional, and national incidence, prevalence, and years lived with disability for 310 diseases and injuries, 1990-2015: a systematic analysis for the Global Burden of Disease Study 2015". Lancet. 388 (10053): 1545–1602<sup>1</sup> The number of patients with drug addiction

Source (Peak Sales): Sales of each indications are extracted form Evaluate Pharma's data of sales by disease and sales by individual products (as of 25 December 2024).<sup>2</sup> Nxera may target one segment in the market for specific diseases



### Partnered pipeline (1/2)

Compound	Target / Mechanism of Action	Modality	Indication	Partner	Disc.	PCC	Ph1	Ph2	Ph3	Арр	Mk
Partnered											
Seebri® Breezhaler®	LAMA	SME	COPD	🔱 novartis							
Ultibro® Breezhaler®	LAMA+LABA	SME	COPD	🐌 novartis							
Enerzair® Breezhaler®	LAMA+LABA+ICS	SME	Asthma	🔥 novartis							
ORAVI®	Antifungal agent miconazole	SME	Oropharyngeal candidiasis	Aisamitsu	_						
Cenerimod	S1P <sub>1</sub> receptor modulator	SME	SLE		_						
NBI-1117568	Muscarinic M4 agonist	SME	Schizophrenia	<b>SIDSCIENCES</b>	_						
NBI-1117568	Muscarinic M4 agonist	SME	Bipolar Mania	<b>SIDSCIENCES</b>							
NBI-1117569	Muscarinic M4 preferring agonist	SME	Neurology diseases	S NEUROCRINE BIOSCIENCES			_				
NBI-1117570	Muscarinic M1/M4 agonist	SME	Neurology diseases	S NEUROCRINE BIOSCIENCES	_		_				
NBI-1117567	Muscarinic M1 preferring agonist	SME	Neurology diseases	Siosciences			_				
PF-07054894	CCR6 antagonist	SME	Inflammatory bowel disease	<b>Pfizer</b>			_				
PF-07258669	MC4 antagonist	SME	Malnutrition	2 Pfizer			_				
PF-06954522	GLP-1 agonist	SME	Type 2 Diabetes	2 Pfizer			_				
(Not disclosed)	CGRP antagonist	SME	Neurology diseases	2 Pfizer							
(Not disclosed)	Multi target	SME/LME	Multiple indications	Genentech A Member of the Roche Group	_						
(Not disclosed)	Multi target	SME	Neurology	abb√ie	_						
(Not disclosed)	Multi target	SME	Diabetes/Metabolic	Lilly	_						

Note: SME = small molecule. LME = large molecule. Seebri<sup>®</sup>, Ultibro<sup>®</sup>, Enerzair<sup>®</sup> and Breezhaler<sup>®</sup> are registered trademarks of Novartis AG.



## Partnered pipeline (2/2)

Compound	Target / Mechanism of Action	Modality	Indication	Partner	Disc.	PCC	Ph1	Ph2	Ph3	Арр	Mkt
Co-development											
KY1051	CXCR4 mAb	mAb	Immuno-oncology	sanofi							
(Not disclosed)	AI-Augmented Drug Discovery	SME	Neurology diseases	, D PHARMENABLE	_						
(Not disclosed)	Multi targe	SME/LME	Immune / Neurology diseases	precisionlife	_						
Co-owned compan	ies										
TMP-301	mGlu5 NAM	SME	Alcohol use disorders					- 1			
TMP-301	mGlu5 NAM	SME	Cocaine use disorders								
ORX750	OX2 agonist (Oral)	SME	Narcolepsy Type 1/2, IH					- 1			
ORX142	OX2 agonist (Oral)	SME	EDS in neurology								
ORX489	OX2 agonist (Oral)	SME	Neurology								



## In-house pipeline

Compound	Target / Mechanism	Modality	Indication	Partner	Disc.	PCC	Ph1	Ph2	Ph3	Арр	Mkt
In-house Programs											
<b>PIVLAZ®</b>	ETA antagonist	SME	Cerebral vasospasm	NXELO 🗸							
QUVIVIQ™	Dual Orexin antagonist	SME	Insomnia	SHIONOGI							
NXE0048149 <sup>1</sup>	GPR52 agonist	SME	Neurology diseases	Boehringer Ingelheim			_				
NXE0039732	EP4 antagonist	SME	Immuno-oncology	NXELCI 🛪			_				
NXE0033744	EP4 agonist	SME	Inflammatory bowel disease	NXEFCI 🛰			_				
NXE0027477	GPR35 agonist	SME	Inflammatory bowel disease	NXELCI 🛪							
(Not disclosed)	Muscarinic M1 agonist (JP)	SME	Neurology diseases	NXELCI 🛪							
(Not disclosed)	SARS CoV-2 Mpro	SME	Coronaviruses	NXELC	_						
Multiple programs	Not disclosed	SME/LME	Neurology diseases	NXELCI 🛪	_						
Multiple programs	Not disclosed	SME/LME	GI and Inflammatory diseases	NXELC	_						
Multiple programs	Not disclosed	SME/LME	Immunology diseases	NXera	_						
In-house Programs (No	longer internally funded. Targeting	g academic / i	ndustrial partnership)								
NXE'310	SSTR5 agonist	Peptide	Hypoglycaemic disorders	NXELQ ~							
NXE'097	GLP-1 antagonist	Peptide	Hypoglycaemic disorders	NXELQ ~							
NXE'023	Dual GLP-2/GLP-1 agonist	Peptide	Intestinal failure/NASH	NXEFO A							
(Not disclosed)	Apelin agonist	Peptide	Pulmonary Arterial Hypertension	NXELQ ~							
NXE'641	Dual orexin antagonist	SME	Insomnia and sleep disorders	NXELCI ~							
(Not disclosed)	PAR-2 mAb	mAb	Atopic Dermatitis/Pain	NXELCI							





#### **Clinical Trials**

Compound	MoA	Condition	Phase	Size	Patient	Start	Completion*	Last Update	Link (main/latest)	Link (others)
NBI-1117568	M4 agonist	Schizophrenia	Ph2	210	Yes	2022-10-04	2024-07-10	2024-09-27	NCT05545111	-
NBI-1117569	M4 preferring agonist	Neurology diseases	Ph1	-	-	-	-	-	-	-
NBI-1117570	M1/M4 agonist	Neurology diseases	Ph1	-	No	2024-03-11	2025-09-04	2024-10-30	2023-508814-40-00	-
NBI-1117567	M1 preferring agonist	Neurology diseases	Ph1	-	-	-	-	-	-	-
PF-07054894	CCR6 antagonist	Inflammatory bowel diseases	Ph1	27	Yes	2022-11-07	2026-01-14	2025-03-25	NCT05549323	NCT06327880 NCT04388878
PF-07258669	MC4 antagonist	Malnutrition	Ph1	14	No	2025-01-02	2025-02-11	2024-11-27	NCT06706869	NCT04628793 NCT05113940
PF-06954522	GLP-1 agonist	T2DM/Obesity	Ph1	45	Yes	2024-02-20	2025-04-07	2025-02-12	NCT06279234	NCT06393517 NCT06003777
TMP-301	mGlu5 NAM	Alcohol use disorders	Ph2	100	Yes	2024-11-14	2025-11-15	2024-12-19	NCT06648655	-
TMP-301	mGlu5 NAM	Cocaine use disorders	Ph1	18	Yes	2025-01-04	2025-08-15	2025-03-25	NCT06648668	-
ORX750	OX2 agonist	Narcolepsy Type 1/2, IH	Ph2	78	Yes	2024-12-23	2025-12	2024-12-31	NCT06752668	-
NXE0048149	GPR52 agonist	Neurology diseases	Ph1	24	No	2024-06-07	2025-11-15	2024-09-23	ISRCTN44913564	ISRCTN17231793
NXE0039732	EP4 antagonist	Immuno-oncology	Ph1/2	150	Yes	2023-07-13	2026-09	2024-12-02	NCT05944237	-
NXE0033744	EP4 agonist	Inflammatory bowel diseases	Ph1	Up to 220	-	2023-11-24	2026-06-30	2024-02-15	ISRCTN70080074	-



#### Exchange Rate, Intangible Assets and Non-core Costs

#### Average exchange rate during period

		FY2025	FY2024	FY2023	FY2022
USD:JPY	Actual	-	151.43	140.53	131.30
	Estimate	152	140	143	
GRP:JPY	Actual	-	193.49	174.81	161.76
	Estimate	193	172	166	

Intangible assets (JPY mi								
	Dec 31, 2024	Dec 31, 2023	Dec 31, 2022					
PIVLAZ®	36,164	37,527	-					
Core technology	8,365	8,466	8,217					
QUVIVIQ™	6,825	5,825	-					
Customer-related assets	227	227	219					
Oravi®	78	89	101					
Other	252	157	40					
Total	51,911	52,291	8,577					

Non-core costs (full year)								
	FY 2024	FY 2023	FY 2022					
Cost of sales adjustment	2,401	1,812	-					
Amortization	2,371	1,495	782					
M&A related costs	1,220	1,263	-					
Depreciation	1,613	983	563					
Share-based Payments	1,396	844	542					
Restructuring costs	28	53	533					
Impairment	-	-	-					
Total	9,029	6,450	2,420					

#### NXera

PIPELINE

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### Glossary

	Basic Terminology/Technology		
GPCR	G Protein-Coupled Receptor	There are about 800 types of GPCRs in the human body. While 400 of them are known to be potential drug targets, about 300 of them are not yet drugged	
NxStaR™	Stabilized Receptor	Nxera' proprietary technology to stabilize a GPCR by engineering a small number of single point mutations outside of the ligand-binding site. It enables to identify the structure of GPCRs to be used for SBDD drug discovery as well as antibody drug discovery as antigens	
SBDD	Structure-Based Drug Design	A method to design drugs on a computer base based on the analysis of the three-dimensional structure of the drug target (e.g., protein receptor)	
TPD	Targeted Protein Degradation	Drugs that promote the degradation of target proteins (e.g., receptors) in cells and aim for therapeutic effects by reducing disease-causing proteins	
PAM	Positive Allosteric Modulator	A regulator that binds to unusual active sites (allosteric sites) on the receptor to increase the affinity and effect of the agonist	
NAM	Negative Allosteric Modulator	A regulator that binds to an unusual active site on the receptor (allosteric site) and reduces the affinity and effectiveness of the agonist	
Ag	Agonist	A therapeutic drug that binds to a receptor and activates an intracellular signaling system similar to biological substances	
Ant	Antagonist	A therapeutic drug that suppresses biological reactions by binding to receptors and preventing them from binding to biological substances	
PK	Pharmacokinetics	Research and testing on the relationship between drug dosage and blood concentration. Mainly describes the rate process of ADME	
PD	Pharmacodynamics	Research and testing on the relationship between drug concentration and pharmacological effects	
ADME	Absorption, Distribution, Metabolism and Excretion	A series of process in the absorption of drugs into the body, distribution within the body, metabolism in the liver and other organs, and excretion in the kidneys and other organs	
POM	Proof of Mechanism	Proof of mechanism of action, mainly through biomarkers. It can suggest the possibility of efficacy in fewer cases than POC	
POC	Proof of Concept	Proof of a therapeutic concept, primarily through clinical efficacy and safety	
Ach	Acetylcholine	A neurotransmitter released from the peripheral parasympathetic and motor nerves to transmit nerve stimuli	
IND	Investigational New Drug	Information packages for development candidates to be submitted to the U.S. Food and Drug Administration (FDA) at the time of initiation of clinical trials	
Ph1	Phase1	A study in humans. The main purpose is to confirm the safety of the drug candidate mainly by healthy volunteers.	
Ph2	Phase2	A study in humans. The main purpose is to confirm the efficacy of the drug candidates on a small scale (however, the number of patients varies greatly depending on the disease)	
Ph3	Phase3	A study in humans. The main purpose is to determine the efficacy of the drug candidates on a large scale (however, the number of patients varies greatly depending on the disease)	
NDA	New Drug Application	An application to the U.S. Food and Drug Administration (FDA) for approval to market a new drug	

Disease/Drug			
LAMA	Long Acting Muscarinic Antagonist	An inhalant that dilates bronchial tubes and improves respiratory function by inhibiting the action of acetylcholine receptors (M3), which increase parasympathetic nerves.	
LABA	Long Acting Beta2-Agonist	An inhalant that improves respiratory function by stimulating sympathetic beta2 receptors to dilate the bronchi.	
ICS	Inhaled Corticosteroid	An inhalant that suppresses airway inflammation to prevent coughing attacks and other symptoms caused by asthma, also promotes the action of beta 2 stimulants and improve airway hyperresponsiveness.	
mCRPC	Metastatic Castration–Resistant Prostate Cancer	Cancer that has spread (metastasized) beyond your prostate gland and for which hormone therapy is no longer effective in stopping or slowing the disease.	
COPD	Chronic Obstructive Pulmonary Disease	A group of diseases that causes damage to the bronchi and lung due to smoking or inhalation of toxic substances, resulting in breathing problems.	
AD	Alzheimer's Disease	Alzheimer's disease is a progressive neurologic disorder that causes the brain to shrink (atrophy) and brain cells to die, the most common cause of dementia.	
DLB	Dementia with Lewy Bodies	Protein deposits, called Lewy bodies, develop in nerve cells in the brain regions involved in thinking, memory and movement (motor control), the second most common type of dementia.	



#### Locations



Midtown East, 9-7-2 Akasaka Minato-ku Tokyo 107-0052

Japan



F17, 410 Teheran-Ro GangHam-Gu Seoul 06192

South Korea



Steinmetz Building Granta Park, Cambridge CB21 6DG

United Kingdom



Spaces Grosspeter Tower, Grosspeteranlage 29, 4052 Basel

Switzerland

# Thank you

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