NXera ~

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

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Agenda

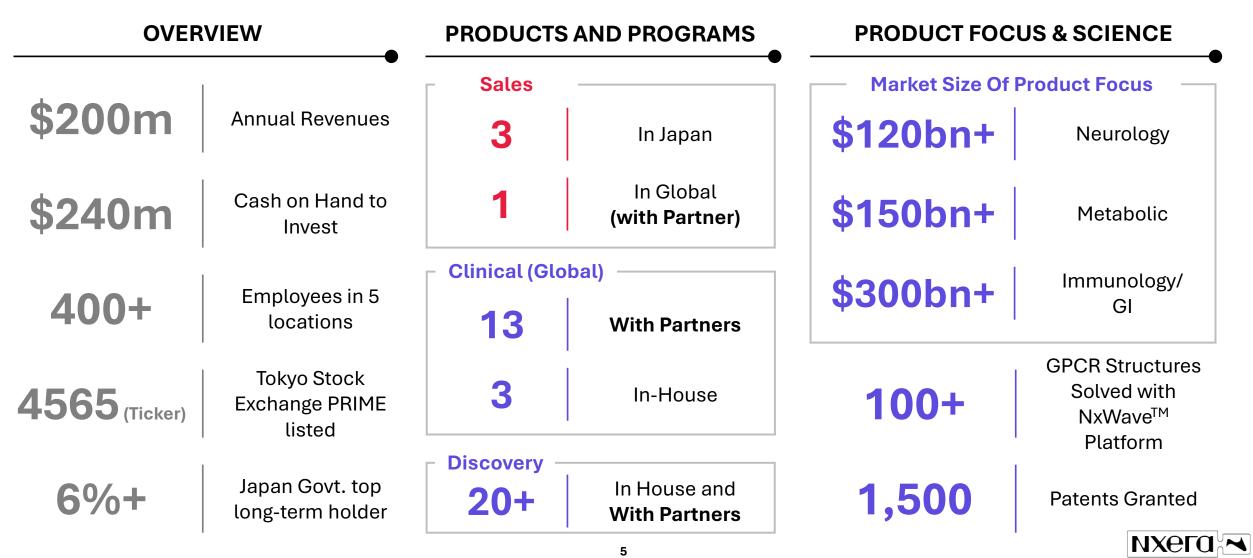
- Business Overview
- Strategic Roadmap
- Our Pipeline
- Japan/APAC Business
- •• Our NxWaveTM Platform
- Financial Results
- Appendix



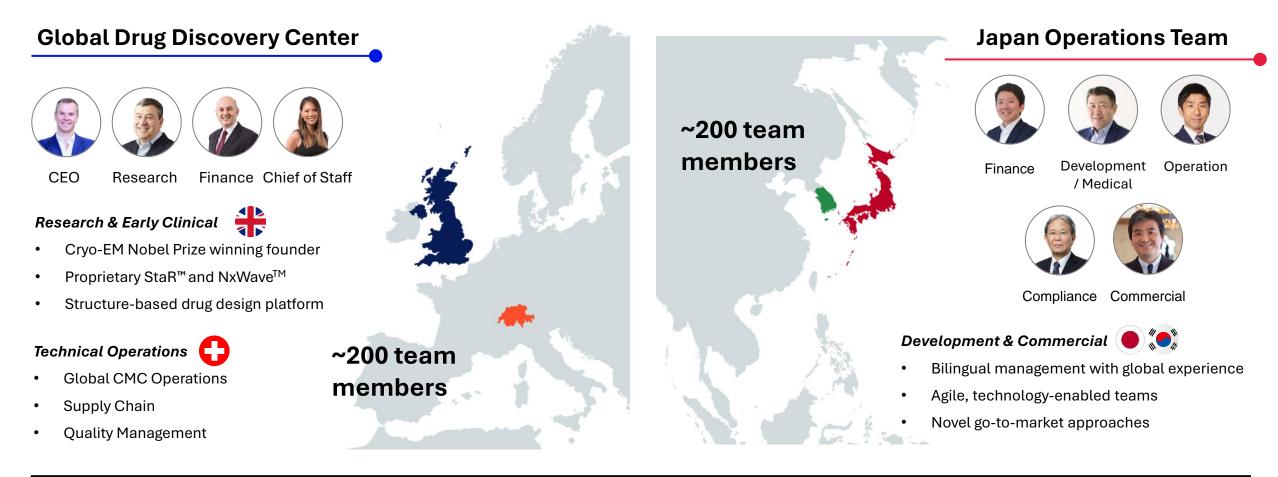
Business Overview

We are Nxera Pharma

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



Not a traditional Japanese pharma. We think and innovate globally, and specialize locally



Our team is committed to addressing some of the biggest healthcare challenges globally



Strategic Roadmap

PIPELINE

JP/APAC

PLATFORM

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

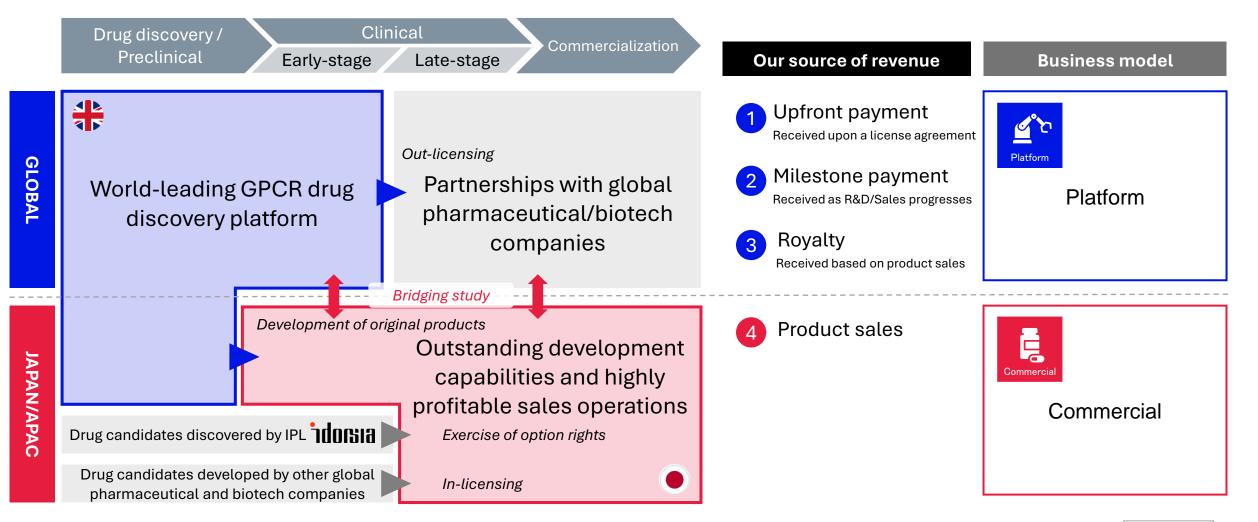
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Building a fully integrated biopharma from Japan

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





PLATFORM FINANCIALS

JP/APAC

Priority objectives for FY2025

01) <u>JPY 17 billion+</u> Net product sales (PIVLAZ[®] plus QUVIVIQ[®])



Commercial

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



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)



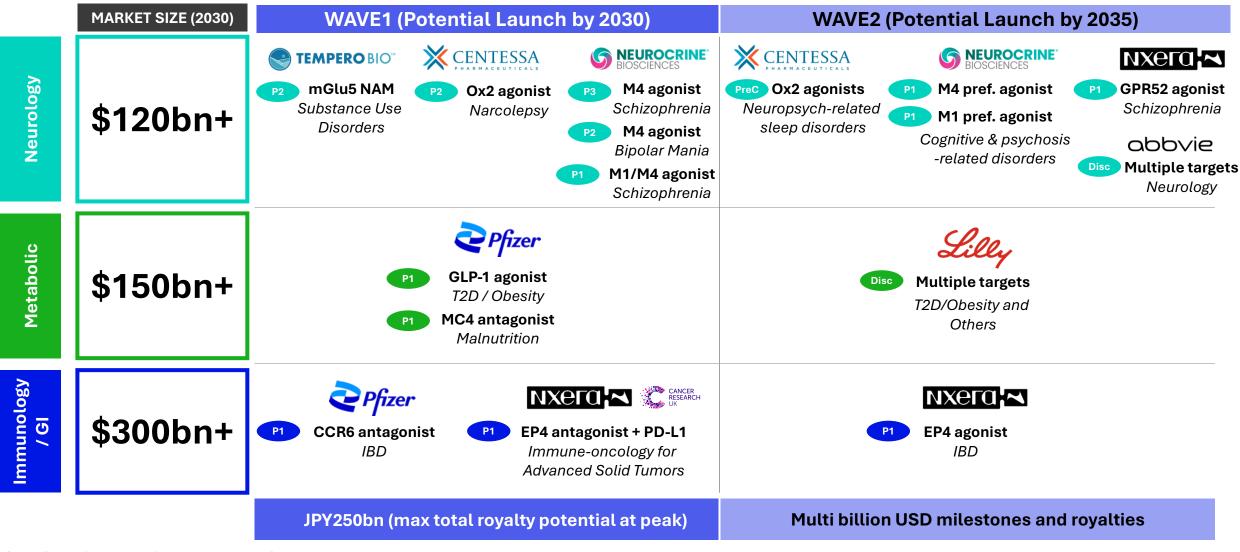


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Partner's Wave 1 and Wave 2 programs are positioned across fast growing disease areas of healthcare



Source: EvaluatePharma, News Research, Internal Analysis

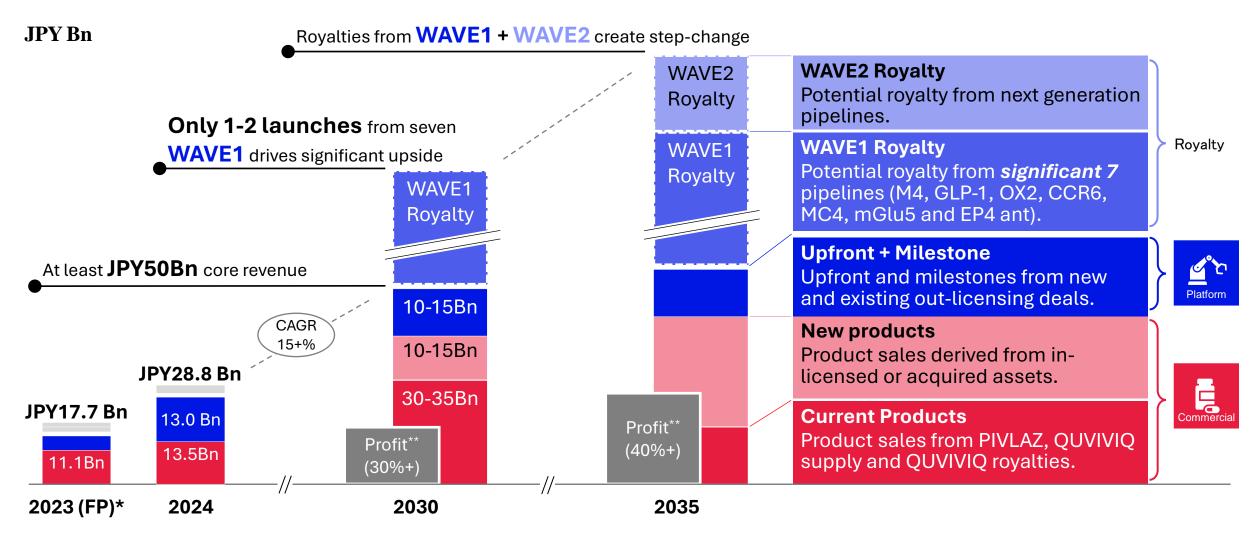
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Our 2030 vision is to build a high growth, highly profitable Japanese biopharma



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.



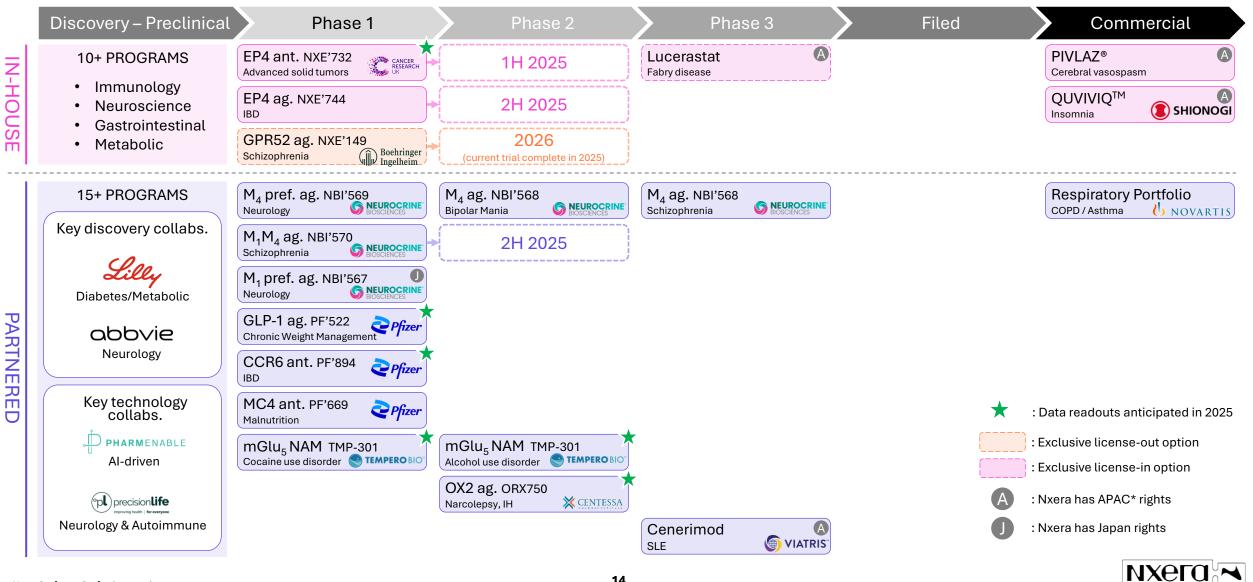
Our Pipeline Programs by Design



JP/APAC PLATFORM FINANCIALS

APPENDIX

Major pipeline Overview (incl. projections)



Note: Pref. ag. : Preferring agonist

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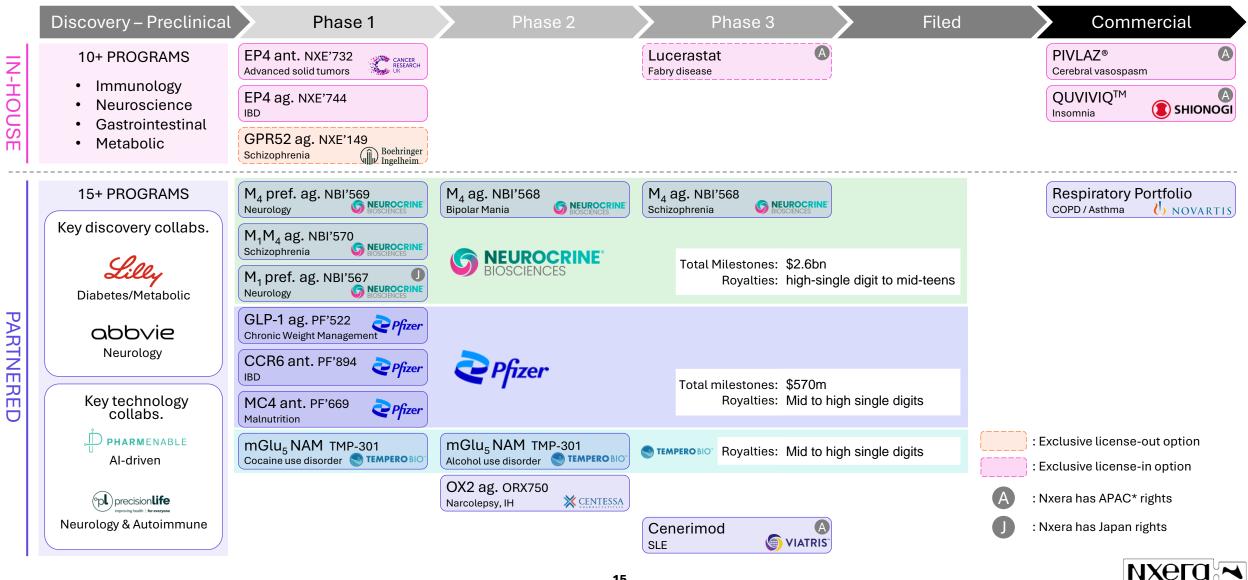
*APAC (ex-China) territory includes Japan, South Korea, Australia, Brunei, Cambodia, Indonesia, Laos, Malaysia, Myanmar, New Zealand, Philippines, Singapore, Taiwan, Thailand and Vietnam

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Major pipeline Overview (incl. key partner highlights)



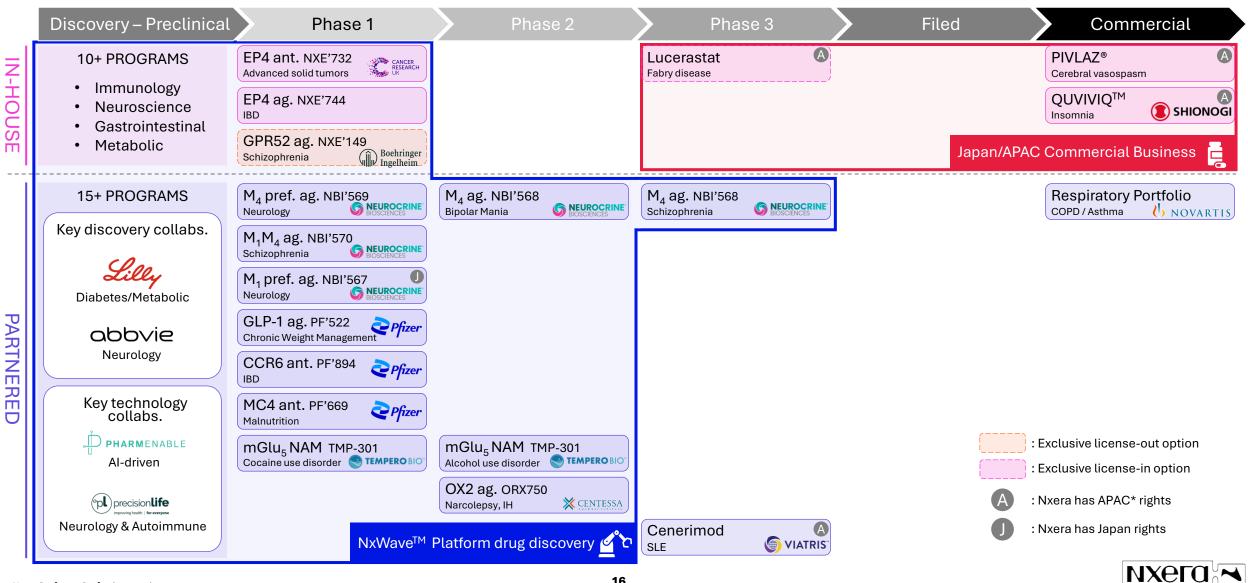
Note: Pref. ag. : Preferring agonist

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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 (By business categories)



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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: Progress in 2025

Looking ahead to potential catalysts in 2025*

Looking undue to potential educitysts in 2025					
PROGRAM	PARTNER	TIMING	EVENT		
Cenerimod		Feb. 2025	Assignment of JAPAC rights		
TMP-301 (mGlu5 NAM)		Mar. 2025	Phase 2 study start in alcohol use disorder		
NBI'568 (M4 agonist)		Apr. 2025	Phase 3 study start in Schizophrenia		
Lucerastat	ndorsia	H1 2025	Exclusive opt-in decision		
NXE'732 (EP4 antagonist)		H1 2025	Phase 2a study start in Advancing Solid Tumours		
NBI'568 (M4 agonist)	NEUROCRINE BIOSCIENCES	H2 2025	Phase 2 study start in Bipolar Mania		
NBI'570 (M1/M4 agonist)		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)	NXEIO M Boehringer Ingelheim	H2 2025	Phase 1b completion		
NXE'732 (EP4 antagonist)		H2 2025	Phase 1b topline data		
ORX750 (OX2 agonist)		H2 2025	Phase 2 data readout (NT1/NT2/IH)		
TMP-301 (mGlu5 NAM)		End 2025	Phase 2 result in alcohol use disorder		
Multiple discovery collaboration progress	abb∨ie <i>Lilly</i>	2025	Progression through discovery stage		
NBI'567 (M1 ago) / NBI'569 (M4 ago) / NBI'570 (M1/M4 ago)	NEUROCRINE BIOSCIENCES	2025	Phase 1 data readout		
QUVIVIQ™	Holling Bio-Pharma Corp.	Feb. 2025	Out licensing in Taiwan		
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		

* Partnered product progress is as already signaled or disclosed by partner



Key strategy for each business category

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

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

STRATEGIC ROADMAP

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OVERVIEW



Japan/APAC Business

Deliver innovation to patients in Japan/APAC

OVERVIEW STRATEGIC ROADMAP PIPELINE JP/APAC PLATFORM

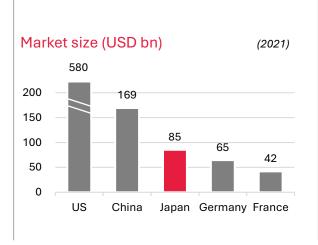


APPENDIX

Japan will serve as our base to expand across APAC markets

Japan is an attractive, established market with strong volumes

Japan is the second largest pharma market (ex-China)



Tailwinds from nearterm regulatory changes

High quality clinical and regulatory environment

APAC is the second highest growth pharma market

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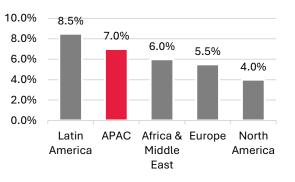
Japan Phase 1 Drug Clinical Trials No Longer Needed for Global Clinical Trials

Excellent access to Doctors/HCPs who evaluate novel drugs

- Typically achieve strong patient uptake
- Reduces drug loss and drug lag for Japan patients







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

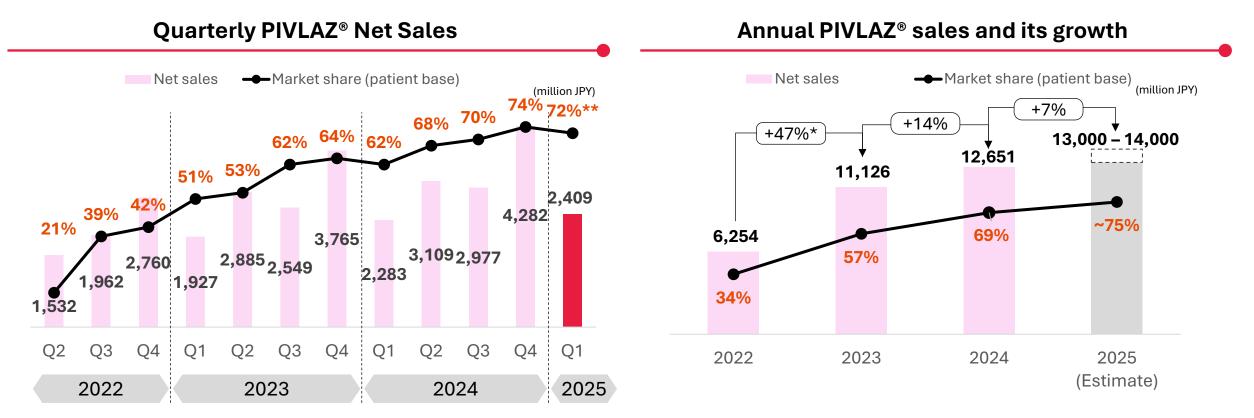


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MHLW

PIVLAZ® (clazosentan, an endothelin A antagonist)

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



STRATEGIC ROADMAP

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エンドセリン受容体拮抗薬

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OVERVIEW

PIVLAZ[®] has rapidly built awareness and is becoming the standard of care with neurosurgeons



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® 点滴静注液

150mg

薬価基準収載

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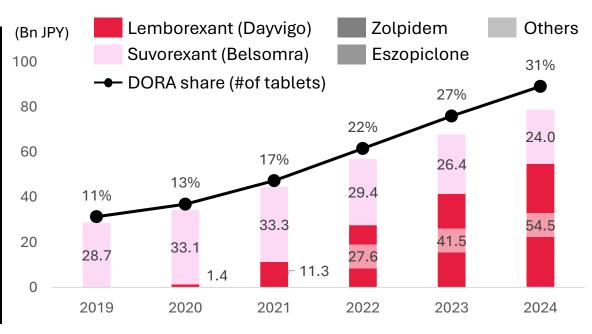
Commercial

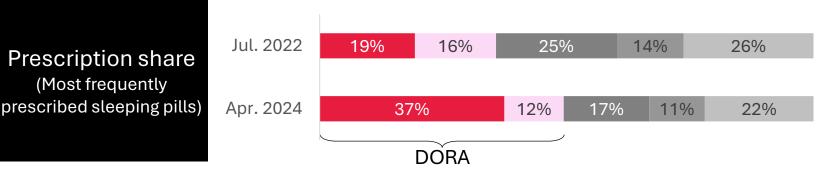
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QUVIVIQTM (daridorexant, dual orexin antagonist "DORA")

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

Sales and market share (NHI-base)







PLATFORM

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- 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

NXero

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OVERVIEW

Quviviq Business structure Image for Nxera

Sales structure

Profit starts from royalty and supply margin will be adding on in a few years



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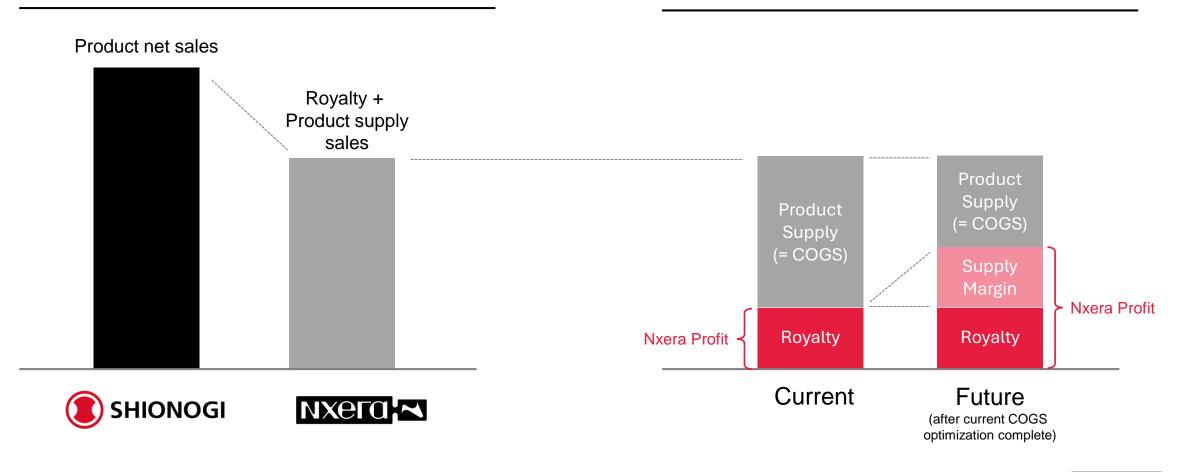
Profit structure for Nxera

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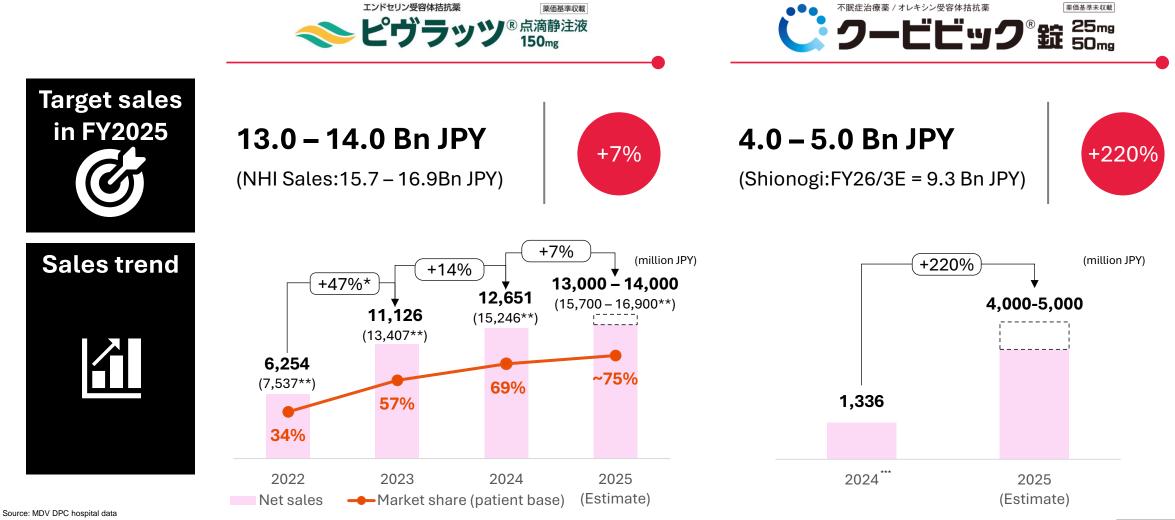




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Full year product sales guidance

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



*: 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 NxWaveTM Platform

Cutting-edge Science

NxWaveTM platform enables faster, cheaper and more precise drug discovery

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

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

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¹ 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.

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

³ 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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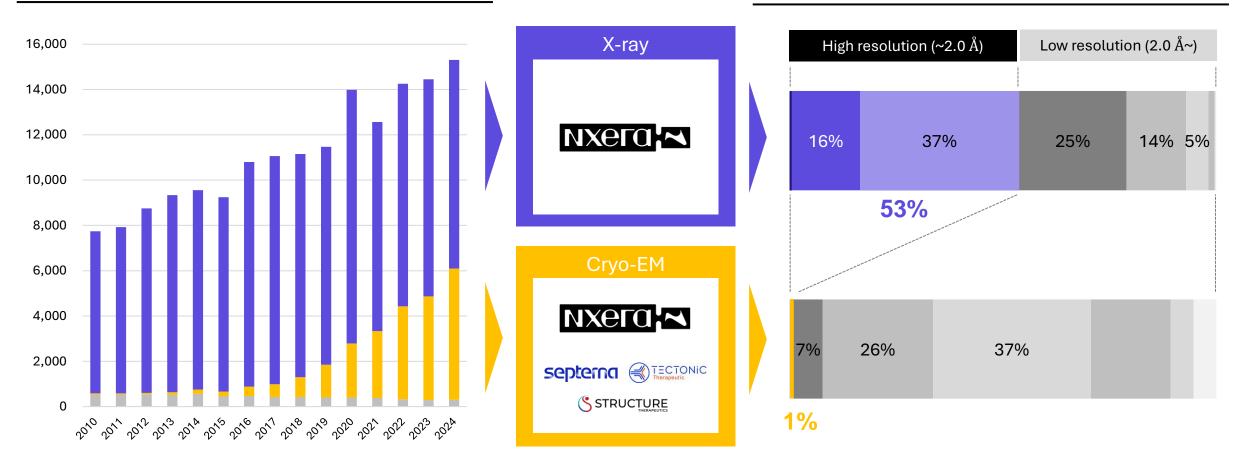
Number of Structures Solved and Deposited in PDB, Resolution by technology

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The number of structures solved using Cryo-EM is increasing, X-ray crystallography has extremely high resolution

Number of stricture solved by technology

Resolution by technology





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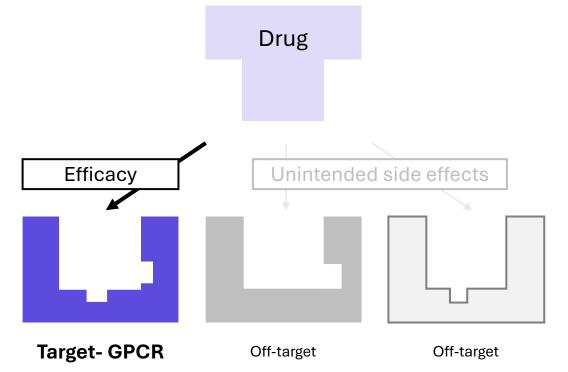


Our platform enables precise design of GPCR models

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

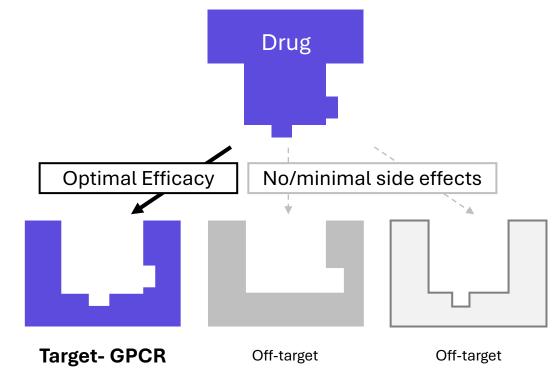
Imprecise GPCR model: Standard Medicine

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

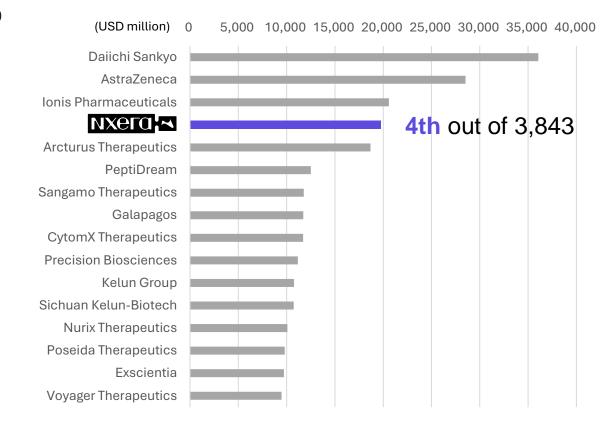
Balance of potential milestone income from existing license agreements¹



Top 15 pharmaceutical/biotech companies by license value² (cumulative total since 2015)

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¹ 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. ² 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)

OVERVIEW

STRATEGIC ROADMAP



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... 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 ¹
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
NEUROCRINE 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
bohaven	December 2020	Collaboration and license agreement for CGRP portfolio	Neurology	\$10m	\$380m
abb√ie	June 2020	Discovery Collaboration and Option to License ²	Inflammatory and Autoimmune	\$32m	\$400m
Takeda	August 2019	Multi-target Collaboration	Multiple; Initial focus on Gastrointestinal	\$26m	\$1.2bn
Genentech A Member of the Roche Group	July 2019	Multi-target Collaboration	Multiple	\$26m	\$1.0bn
P fizer	November 2015	Multi-target Collaboration	Multiple	-	\$1.8bn

¹Potential 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. ² AbbVie has the option to expand the collaboration by an additional three targets



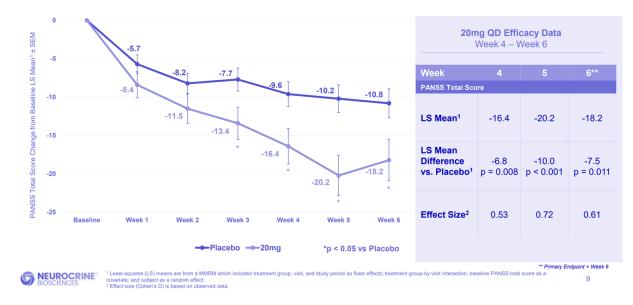


Topline Results for Phase 2 Trial of M4 Agonist 20mg dose demonstrated statistically significant efficacy at Week 3, 4, 5 and 6 vs. placebo

Once-Daily 20mg Dose Met Primary Endpoint PANSS Total Score vs Placebo

Week 6	Placebo N=68	20mg QD N=35	40mg QD N=38	60mg QD N=34	30mg BID N=26
PANSS Total Score					
LS Mean Change from Baseline*	-10.8	-18.2	-12.6	-13.7	-15.8
LS Mean Difference vs. Placebo, p-value*		-7.5 p = 0.011	-1.9 p = 0.282	-2.9 p = 0.189	-5.0 p = 0.090
Effect Size**		0.61	0.27	0.39	0.23

Once-Daily 20mg Dose Demonstrated Clinically Meaningful and Statistically Significant Efficacy at Week 3, 4, 5, and 6



"Least-squares (LS) means are from a MMRM which includes treatment group, visit, and study period as fixed effects; treatment group-by-visit interaction; baseline PANSS total score as a covariate; and subject as a random effect. "Effect size (Cohen 5) is based on observed data.

"The effects with the 20-milligram dose, you see statistical significance between Week 3, 4, 5, and six, meaning that you are seeing a reproducible response here."

Source: Neurocrine presentation - Topline Results for Phase 2 Trial of NBI-1117568 (NBI-'568) in Schizophrenia, August 28, 2024



NEUROCRINE BIOSCIENCES



Mentioned in a presentation Phase 3 of NBI-568 will be one to one randomization and around 20 sites per study

Comparison of Study Sites and Duration with Known Muscarinic Programs

	Neurocrine/Nxera	Neurocrine/Nxera	BMS/Karuna	AbbVie/Cerevel
Compound	NBI-1117568	NBI-1117568	Cobenfy/Kar-XT	CVL-231/Emraclidine
Study name	NCT05545111	NCT06963034	EMERGENT-2/3	EMPOWER-2/3
Route of Administration	oral (once daily)	oral (once daily)	oral (twice daily)	oral (once daily)
Size	213	280+	Total 518	Total 752
Randomization	drug:placebo = 2:1	drug:placebo = 1:1	drug:placebo = 1:1	drug:placebo = 2:1
Number of study sites	15 sites	20 sites (estimate)	22 sites (EMERGENT-2) 32 sites (EMERGENT-3)	26 sites (EMPOWER-2) 25 sites (EMPOWER-3)
Duration	1.8years	2025/5-2027/10(2.2years)	1.6years	2.2years
Phase	Ph2(completed)	Ph3(on trial)	Ph3(completed)	Ph2(unsuccessful)
Primary endpoint	PANSS Total Score Change (Week 6)	PANSS Total Score Change (Week 5)	PANSS Total Score Change (Week 5)	PANSS Total Score Change (Week 5)

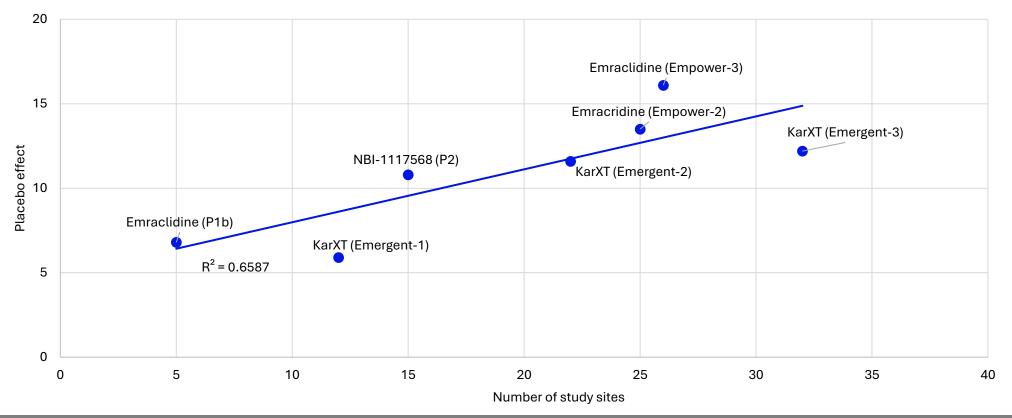


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NEUROCRINE BIOSCIENCES

Data comparison of placebo effects (Total PANSS)

Large number of study sites in a clinical trial of muscarinic program may be linked to a higher placebo response.



"Number of facilities is another important factor in managing the placebo effect"

Source: Neurocrine presentation – Topline Results for Phase 2 Trial of NBI-1117568 (NBI-'568) in Schizophrenia, August 28, 2024



Cobenfy

NEUROCRINE BIOSCIENCES

PLATFORM



Safety: Adverse Events Risk

The gastrointestinal and cardiovascular adverse events were higher than placebo in KarXT, but not on NBI-568

		Placebo N=70	20mg QD N=40	40mg QD N=39	60mg QD N=34	30mg BID N=27	All Treated N=140
	Somnolence	2 (2.9)	5 (12.5)	2 (5.1)	7 (20.6)	1 (3.7)	15 (10.7)
NBI-568	Dizziness	1 (1.4)	5 (12.5)	3 (7.7)	4 (11.8)	1 (3.7)	13 (9.3)
	Headache	14 (20.0)	1 (2.5)	5 (12.8)	1 (2.9)	5 (18.5)	12 (8.6)
	🗙 Nausea	2 (2.9)	2 (5.0)	3 (7.7)	3 (8.8)	0	8 (5.7)
	Constipation	2 (2.9)	2 (5.0)	3 (7.7)	1 (2.9)	1 (3.7)	7 (5.0)

Gastrointestinal (M2)	Cardiovascular (M3)	Others
Similar to placebo	Similar to placebo	Somnolence Dizziness
x3-5 vs. placebo (Four items with 10% or more)	x4 vs. placebo (Occurred in 5.9%)	Dry mouth

Table 3.6. Pooled Treatment-Related Adverse Events in EMERGENT trials²⁰

Adverse Event, %	KarXT (n= 340)	Placebo (n= 343)
★ Nausea	17.1%	3.2%
★ Constipation	15.0%	5.2%
★ Dyspepsia	12.1%	2.3%
★ Vomiting	10.9%	0.9%
★ Hypertension	5.9%	1.2%
Dry Mouth	5.0%	1.5%
Tachycardia	4.7%	2.0%

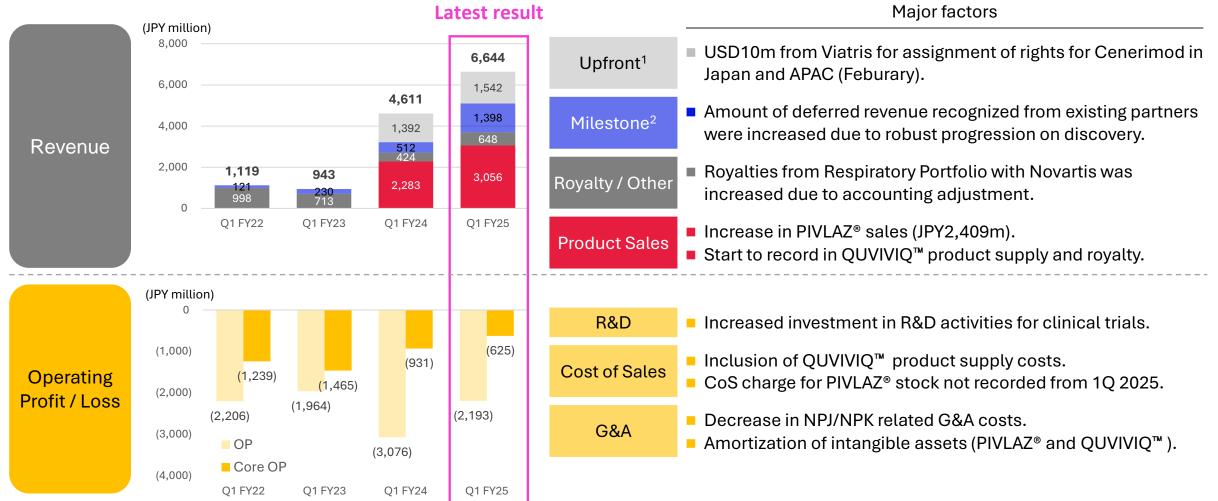
Source: Neurocrine presentation – Topline Results for Phase 2 Trial of NBI-1117568 (NBI-'568) in Schizophrenia, August 28, 2024, KarXT for Schizophrenia draft evidence report Nov. 28, 2023 (https://icer.org/wp-content/uploads/2023/07/ICER_Schizophrenia_Draft_Report_For-Publication_112823.pdf)



Financial Results

Key financial indicators

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



¹ Upfront fee revenue recognised at deal inception

² Milestone revenue recognised at milestone event + deferred revenue releases

Breakdown of Q1 results

Business is progressing smoothly. Significant improvement in revenue from commercial

(JPY million)	<mark>ه ک</mark> Platform ^{*1}		Commercia	l ^{*2}	Consolidate P&L (Core)		Non-core costs	Consolidate P&L (IFRS)	d
Revenue	2,046	(YoY) -12%	4,598	(YoY) +101%	6,644	(YoY) +44%	Total : 1,568	6,644	(YoY) +44%
Cost of Sales	631	+332%	968	+191%	1,599	+234%		1,615	+36%
SG&A	1,189	+17%	1,296	-13%	2,485	-1%	A Amortization (447) Other (785)	3,701	+1%
R&D	3,178	+27%	294	-21%	3,472	+21%	B Other (336)	3,808	+20%
Other income	293	-24	(6)	-6	287	-30	·	287	-30
OP/Core OP	(2,659)	-1,637	2,034	+1,943	Core OP (625)	+306		OP (2,193)	+883

A mortization of intangible assets (currently relates to PIVLAZ[®] and QUVIVIQ^M).

B Amortization of other intangible assets (e.g. IP), depreciation (e.g. laboratory equipment), share-based payments and other restructuring costs.

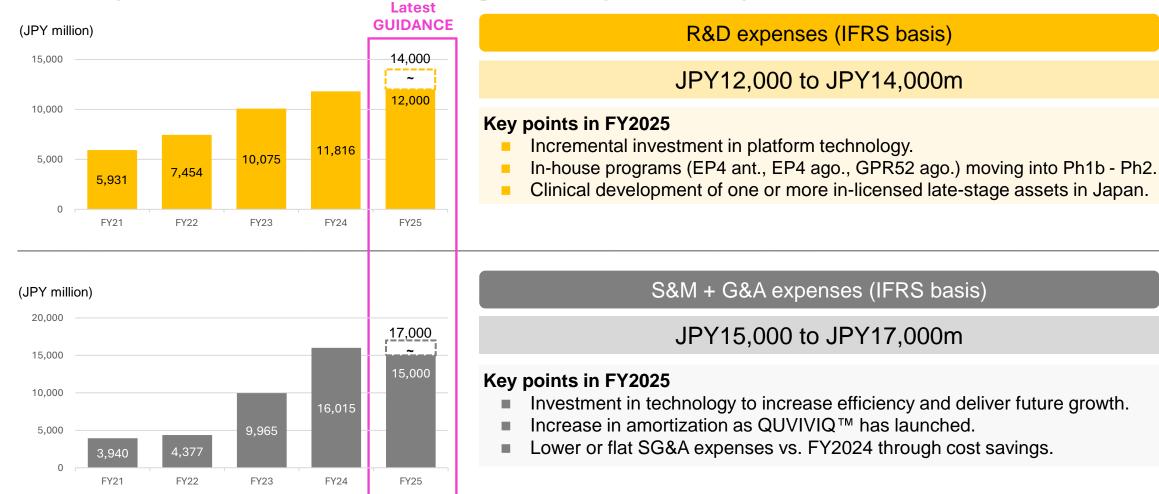
*1 = Nxera Pharma Co. Ltd. (formerly Sosei Group Corporation) + Nxera Pharma UK Ltd (formerly Heptares Therapeutics Ltd.) + Sosei K.K (ex - Nxera Pharma Basel blanch)

*2 = Nxera Pharma Japan (formerly Idorsia Pharmaceuticals Japan) + Nxera Pharma Korea (formerly Idorsia Pharmaceuticals Korea) + Nxera Pharma Basel blanch



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





APPENDIX



Partnered pipeline (1/2)

Compound	Target / Mechanism of Action	Modality	Indication	Partner	Disc.	PCC	Ph1	Ph2	Ph3	Арр	Mkt
Partnered											
Seebri® Breezhaler®	LAMA	SME	COPD	U NOVARTIS							
Ultibro® Breezhaler®	LAMA+LABA	SME	COPD	U NOVARTIS							
Enerzair® Breezhaler®	LAMA+LABA+ICS	SME	Asthma	U NOVARTIS							
ORAVI®	Antifungal agent miconazole	SME	Oropharyngeal candidiasis								
Cenerimod	S1P ₁ receptor modulator	SME	SLE								
NBI-1117568	Muscarinic M4 agonist	SME	Schizophrenia	NEUROCRINE BIOSCIENCES							
NBI-1117568	Muscarinic M4 agonist	SME	Bipolar Mania	NEUROCRINE BIOSCIENCES							
NBI-1117569	Muscarinic M4 preferring agonist	SME	Neurology diseases	NEUROCRINE BIOSCIENCES			_				
NBI-1117570	Muscarinic M1/M4 agonist	SME	Neurology diseases	NEUROCRINE BIOSCIENCES			_				
NBI-1117567	Muscarinic M1 preferring agonist	SME	Neurology diseases	NEUROCRINE BIOSCIENCES			_				
PF-07054894	CCR6 antagonist	SME	Inflammatory bowel disease	Pfizer			_				
PF-07258669	MC4 antagonist	SME	Malnutrition	Pfizer			_				
PF-06954522	GLP-1 agonist	SME	Chronic Weight Management	Pfizer			_				
(Not disclosed)	CGRP antagonist	SME	Neurology diseases	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*, Ultibro*, Enerzair* and Breezhaler* are registered trademarks of Novartis AG.



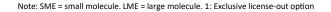
Partnered pipeline (2/2)

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	. PHARMENABLE	_						
(Not disclosed)	Multi targe	SME/LME	Immune / Neurology diseases	precisionlife	_						
Co-owned compan	ies										
TMP-301	mGlu5 NAM	SME	Alcohol use disorder					_			
TMP-301	mGlu5 NAM	SME	Cocaine use disorder								
ORX750	OX2 agonist (Oral)	SME	Narcolepsy Type 1/2, IH					_			
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											
PIVLAZ ®	ETA antagonist	SME	Cerebral vasospasm	NXEFO ~							
QUVIVIQ™	Dual Orexin antagonist	SME	Insomnia	() SHIONOGI							
NXE0048149 ¹	GPR52 agonist	SME	Neurology diseases	Boehringer Ingelheim			_				
NXE0039732	EP4 antagonist	SME	Immuno-oncology	NXELC -			_				
NXE0033744	EP4 agonist	SME	Inflammatory bowel disease	NXELC .~			_				
NXE0027477	GPR35 agonist	SME	Inflammatory bowel disease	NXELC .							
(Not disclosed)	Muscarinic M1 agonist (JP)	SME	Neurology diseases	NXELCI 🛪							
(Not disclosed)	SARS CoV-2 Mpro	SME	Coronaviruses	NXELCI 🛪	_						
Multiple programs	Not disclosed	SME/LME	Neurology diseases	NXELC! ~	_						
Multiple programs	Not disclosed	SME/LME	GI and Inflammatory diseases	NXELC!~	_						
Multiple programs	Not disclosed	SME/LME	Immunology diseases	NXELC	_						
In-house Programs (No	longer internally funded. Targeting	g academic / i	ndustrial partnership)								
NXE'310	SSTR5 agonist	Peptide	Hypoglycaemic disorders	NXELCI 🛃							
NXE'097	GLP-1 antagonist	Peptide	Hypoglycaemic disorders	NXELC							
NXE'023	Dual GLP-2/GLP-1 agonist	Peptide	Intestinal failure/NASH	NXELO 🗸							
(Not disclosed)	Apelin agonist	Peptide	Pulmonary Arterial Hypertension	NXELCI 🛪							
NXE'641	Dual orexin antagonist	SME	Insomnia and sleep disorders	NXELCI 🛪							
(Not disclosed)	PAR-2 mAb	mAb	Atopic Dermatitis/Pain	NXELO							





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	2025-03-14	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	26	No	2024-12-11	2025-02-20	2025-03-07	NCT06706869	NCT04628793 NCT05113940
PF-06954522	GLP-1 agonist	T2DM/Obesity	Ph1	45	Yes	2024-02-20	2025-04-07	2025-02-13	NCT06279234	NCT06393517 NCT06003777
TMP-301	mGlu5 NAM	Alcohol use disorder	Ph2	100	Yes	2024-11-14	2025-11-15	2025-02-21	NCT06648655	-
TMP-301	mGlu5 NAM	Cocaine use disorder	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	2024-04-25	NCT06752668	-
NXE0048149	GPR52 agonist	Neurology diseases	Ph1	24	No	2024-06-07	2025-11-15	2024-11-05	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	2024-05-02	ISRCTN70080074	-



Estimation of potential market size

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

Cotogony	Category Indication ²		Pe	eak Sales	Candidates	
Category		Number of Patients	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 ¹			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 ¹ 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).² Nxera may target one segment in the market for specific diseases



FINANCIALS

APPENDIX

Exclusive Opt-in Rights And ROFN/ROFR¹

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) ²
/ROFN ¹	IDOR-1117-2520	Undisclosed	Immune-mediated disorders	Phase 1*	
	ACT-777991	CXCR3 antagonist	Recent-onset Type 1 diabetes	Phase 1*	

¹ ROFN/ROFR - Right of first negotiation / Right of first refusal

² 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 - Definition

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

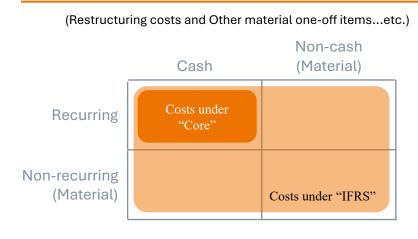
PIPELINE

JP/APAC

OVERVIEW

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

+ Material Non-recurring Costs



Operating Profit **"IFRS"**

PLATFORM

FINANCIALS

APPENDIX

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



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 mn)
	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 yea	r)		(JPY mn)
	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

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
РК	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.				

FINANCIALS

APPENDIX

PLATFORM

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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



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