



37th Annual J.P. Morgan Healthcare Conference 2019

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References to “FY” in this presentation for periods prior to 1 January 2018 are to the 12-month periods commencing in each case on April 1 of the year indicated and ending on March 31 of the following year, and the 9 month period from April 1 2017 to December 31 2017. From January 1 2018 the Company changed its fiscal year to the 12-month period commencing in each case on January 1. References to “FY” in this presentation should be construed accordingly.

About Sosei Heptares (Tokyo Stock Exchange: 4565)

SOSEI FOUNDED 1990, TRANSFORMATIONAL ACQUISITION OF UK BIOTECH HEPTARES 2015

29 PROGRAMS IN R&D TODAY

- 1 in Phase 3
- 2 in Phase 2¹
- 4 in Phase 1²
- 4 in Preclinical Development
- 15 in Discovery

PARTNERSHIPS WITH BIOPHARMA AND ACADEMIA

CURRENT PARTNERS



¹AZD4635 for multiple solid malignancies, HTL0018318 for dementia with Lewy bodies (voluntarily suspended)

²AZD4635 for EGFRm NSCLC, HTL0016878 for neurobehavioral symptoms of Alzheimer's disease, HTL0018318 for Alzheimer's disease (voluntarily suspended), HTL0014242 for neurological disorders

We are a Japan-anchored, integrated global biotech company

R&D CENTER
CAMBRIDGE, UK

~120 EMPLOYEES



HEADQUARTERS
TOKYO, JAPAN

~30 EMPLOYEES



Proprietary StaR^{®1} GPCR technology underpin



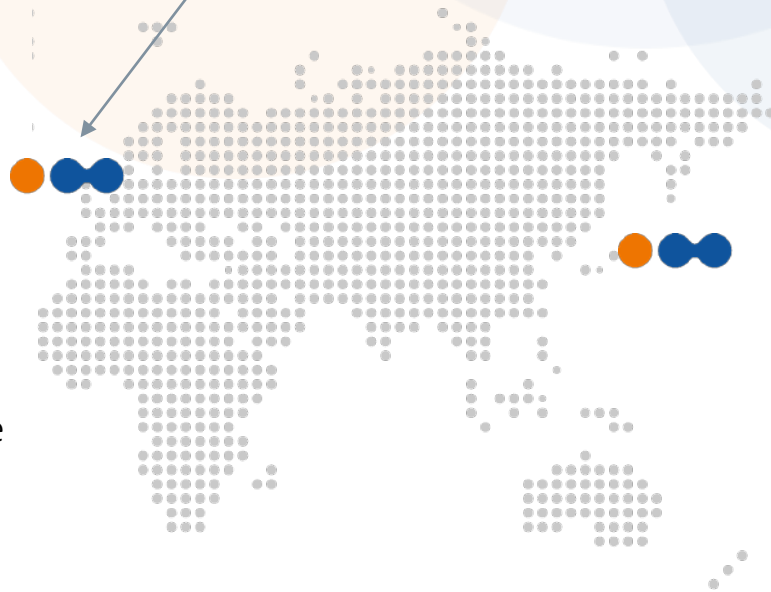
Research, Drug Discovery and SBDD² Platform



Translational and Early-Stage Clinical Development Expertise



Business Development Center



Late-Stage Japanese Development Expertise



Access to Capital and Royalty Income from Novartis

We recently moved to Granta Park, Cambridge - one of the world's top biotech innovation hubs.
Driving enhanced science, productivity, and collaboration and partnership opportunities

¹ Stabilized receptor technology

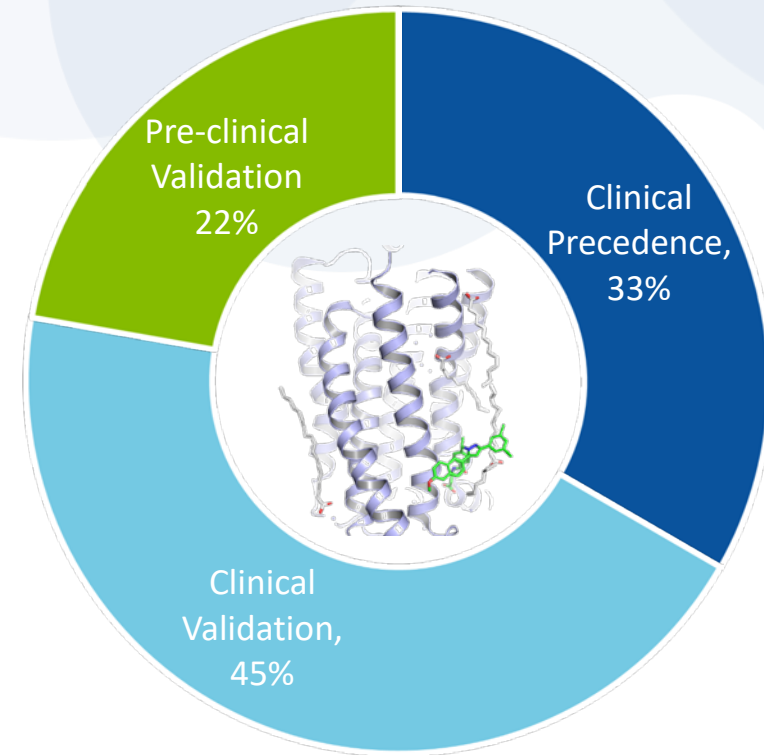
² Structure-based drug design

Our StaR[®] technology enables smarter GPCR drug development

Unique structural insights into GPCRs enable better and smarter drug design

- ✓ Improved physiochemical properties
- ✓ Better safety and efficacy
- ✓ Reduced clinical attrition
- ✓ Small molecule, peptide or antibody discovery

CURRENTLY IN DEVELOPMENT



Unique, scalable and sustainable platform, delivering differentiated pipeline candidates.
Focused on targets with high level of validation

Remaining true to our philosophy as we evolve

- ✓ Technology-driven
- ✓ Keep it small
- ✓ Get published
- ✓ Focus on value creation
- ✓ Be productive



Growth and productivity snapshot

Consistently delivering clinical candidates, PCCs and pre-PCCs

1

We have **identified 22 agents in total vs. 18 targets**, 12 of which were identified since 2015

- **c.2.5** per annum on average (2010–2018)
- **4** clinical agents (3 targets)
- **8** PCCs (Nominated Candidates, 7 targets)
- **6** pre-PCCs (Candidate Selection stage)
- **2** mAbs in Candidate Selection
- **2** pre-PCCs in Pfizer Collaboration

2

Projects take on average **c.2.5 years to identify a PCC** and this has **reduced to c.2.0 years** since 2015

- The PCC is now generally synthesized in **<500 compounds**

Productivity significantly exceeds industry averages

PCC = preclinical candidate, mAbs = monoclonal antibodies

Our model is designed to create and capture optimal value

Reserving the right to choose the best strategy for our proprietary assets

Business model strategy

Out-licensing

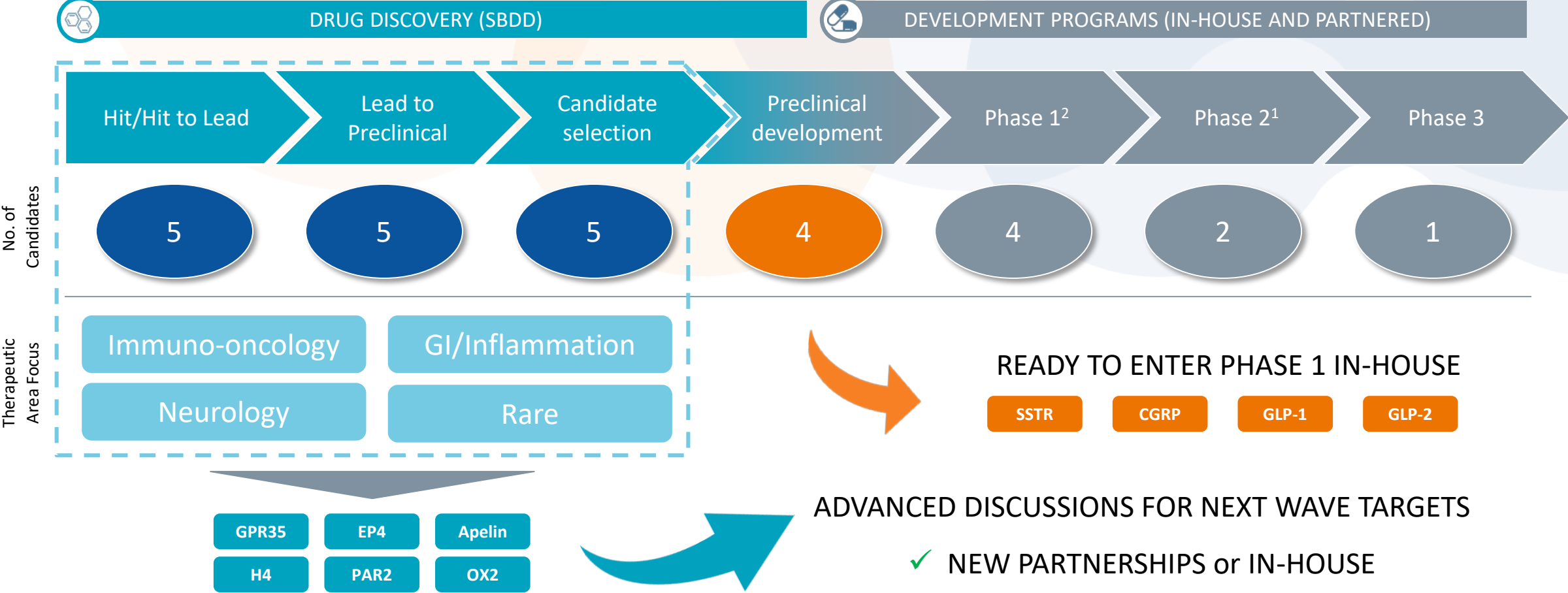
Co-develop / joint venture

In-house discovery and development

Characteristics

- Very large indications
 - Significant clinical trial costs involved
 - Late-stage development expertise required
 - Scale commercialization effort needed
-
- Complementary technology platform (Peptidream – PDPS peptides, Kymab – mAbs)
 - Ability to accelerate program & value creation via creation of asset-centric vehicle
-
- Rare/orphan/specialty disease setting
 - Peak sales opportunity \$300m+
 - Manageable translation and clinical pathway, and digestible development costs

Next wave of novel candidates ready to create value in 2019



Aggressively pursuing new partners for multiple discovery and development deals

¹AZD4635 for multiple solid malignancies, HTL0018318 for dementia with Lewy bodies (voluntarily suspended)
²AZD4635 for EGFRm NSCLC, HTL0016878 for neurobehavioral symptoms of Alzheimer’s disease, HTL0018318 for Alzheimer’s disease (voluntarily suspended), HTL0014242 for neurological disorders

AZD4635: announcing a new \$15m milestone from AstraZeneca

Reverse local immune suppression by blockade of Adenosine 2a Receptor pathway

Partnered with:
AstraZeneca

AZD4635
(A2aR)

Demonstrated progress with AstraZeneca in I/O

- ✓ \$10m upfront (2015)
- ✓ \$10m milestone Phase 1 (2016)
- ✓ \$12m synergy milestone (2017)
- ✓ **\$15m milestone (2019)**

Next-generation I/O therapies to enhance treatment

AZD4635

A2aR antagonist

MONOTHERAPY

AZD4635

A2aR antagonist



durvalumab

Anti-PD-L1

AZD4635

A2aR antagonist




oleclumab

Anti-CD73

AZD4635 emerging as a next-generation I/O therapy that may enhance and broaden efficacy of approved checkpoint inhibitors across more tumor types

AstraZeneca testing AZD4635 in Phase 1b/2 studies

AZD4635 as monotherapy or in combination in tumors of high unmet need

Partnered with:
AstraZeneca 

AZD4635
(A2aR)

ClinicalTrials.gov Identifier: NCT02740985

**Monotherapy
AZD4635
(A2aR antagonist)**

- I-O naïve and post immunotherapy tumors

AZD4635
Post IO NSCLC

AZD4635
IO naïve mCRPC

AZD4635
IO naïve CRC

AZD4635
Other IO naïve

AZD4635
Post IO other

Primary
completion date
2020

**Combo with
durvalumab
(anti-PD-L1)**

- I-O naïve and post immunotherapy tumors

AZD4635 + Durvalumab
Post IO NSCLC

AZD4635 + Durvalumab
IO naïve mCRPC

Primary
completion date
2020

ClinicalTrials.gov Identifier: NCT03381274

**Combo with
oleclumab
(anti-CD73)**

- Locally advanced/metastatic NSCLC with EGFR mutation

AZD4635 + Oleclumab
NSCLC with EGFRmut

Primary
completion date
2021

Partnered pipeline

Multiple shots on goal with world-leading partners across areas of high-unmet need

Product/Program	Modality ¹	Indication	Partner	Discovery	Preclinical	Phase 1	Phase 2	Phase 3	Marketed
Partnered Pipeline - Legacy Respiratory Products (Traditional out-licensing)									
Seebri®/Ultibro®	SME	COPD	NOVARTIS	[Progress bar from Discovery to Marketed]					
QVM149	SME	COPD	NOVARTIS	[Progress bar from Discovery to Phase 3]					
Partnered GPCR Pipeline (Traditional out-licensing/collaboration projects)									
A2a	SME	Multiple solid tumors	AstraZeneca	[Progress bar from Discovery to Phase 2, next 12-18 months progress]					
A2a	SME	EGFRm NSCLC	AstraZeneca	[Progress bar from Discovery to Phase 1]					
M ₁	SME	Alzheimer's disease	Allergan	[Progress bar from Discovery to Phase 1]					
M ₄	SME	Alzheimer's disease	Allergan	[Progress bar from Discovery to Phase 1, next 12-18 months progress]					
M ₁ /M ₄ dual	SME	Alzheimer's disease	Allergan	[Progress bar from Discovery to Phase 1]					
Multiple targets	SME	Pain	Daiichi-Sankyo	[Progress bar from Discovery to Phase 2, next 12-18 months progress]					
Multiple targets	SME/mAb	Multiple indications	Pfizer	[Progress bar from Discovery to Phase 2, next 12-18 months progress]					
Multiple targets	mAb	Inflammation	morphosys	[Progress bar from Discovery to Phase 2, next 12-18 months progress]					
Partnered GPCR Pipeline (Co-development/profit share)									
Multiple targets	mAb	Immuno-oncology	kymab	[Progress bar from Discovery to Phase 2, next 12-18 months progress]					
Multiple targets	Peptide	Inflammation	PeptiDream	[Progress bar from Discovery to Phase 1]					









—● : Current stage
 - -◆ : Next 12–18 months progress

Multiple big pharma partners, across multiple modalities, validate our StaR® and SBDD approach

¹ Note: SME = small molecule; mAb = monoclonal antibody

In-house pipeline

Rapidly emerging pipeline focused on rare/orphan/specialty disease categories

Product/Program	Modality ¹	Indication	Originator	Discovery	Preclinical	Phase 1	Phase 2	Phase 3	Marketed	
Proprietary GPCR Pipeline (Go-to-market/commercialize)										
M ₁	SME	DLB (Japan)		—●—						
mGluR ₅	SME	Neurology		—●—						First Patient Dosed in Ph 1 December 2018 
SSTR	Peptide	Endocrine disorders		—●—◆						UK regulatory authority and ethics committee approval received. First Patient Dosing expected Q1 2019 
CGRP	SME	Migraine		—●—◆						
GLP-1	Peptide	Metabolic diseases		—●—◆						
GLP-2	Peptide	Intestinal failure		—●—◆						

Multiple candidates entering clinical development

—●— : Current stage
 - - -◆ : Next 12–18 months progress

¹ Note: SME = small molecule

Frequently asked questions

1

Has the strategy changed with a new CEO?

No change, strategy remains as outlined at November 2018 results

2

Are there any updates on M₁ tox issue?

Investigative work with Allergan progressing well

3

Is the advent of Cryo-EM impacting your competitive advantage?

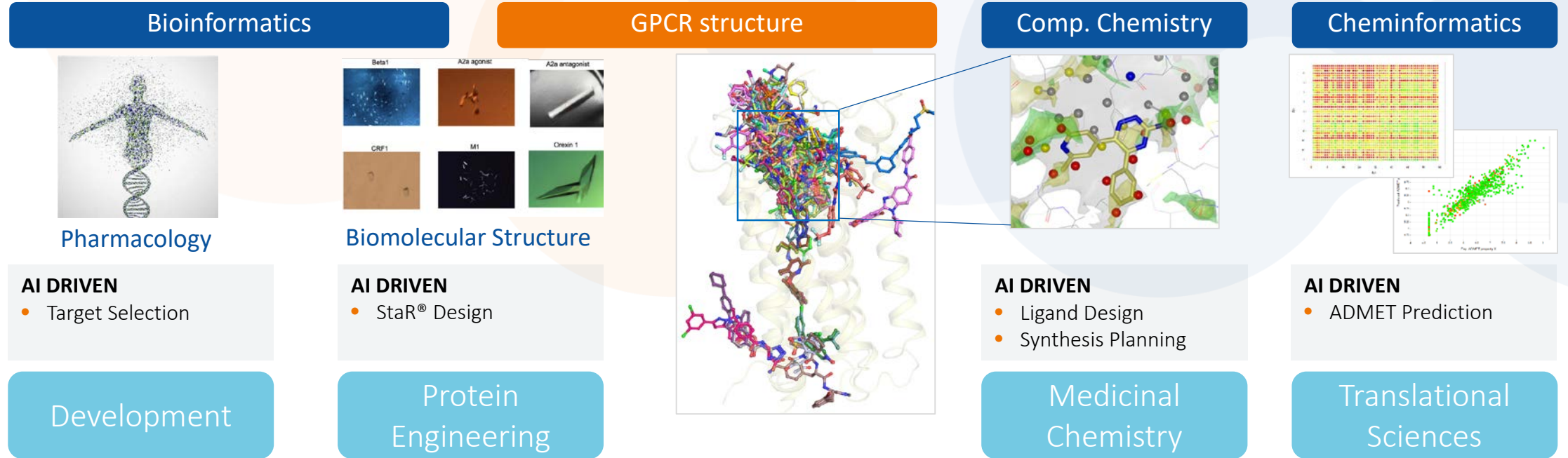
No, Cryo-EM is certainly revolutionary but it will not impact our competitive advantage

4

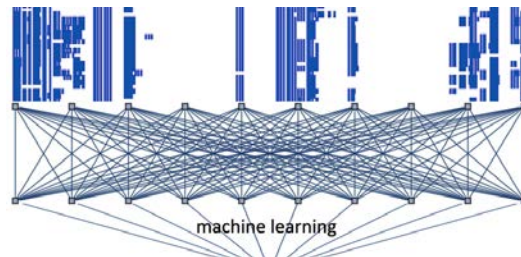
Is there a role for AI / machine learning for GPCRs?

Yes, definitely – we are collaborating with Univ. of Cambridge to deploy AI / machine learning

We are rolling out AI across our drug discovery platform



Data & descriptors
Machine Learning



Artificial Intelligence for Multi-Parametric GPCR Drug Discovery

Thank you!



VISION

To become a leading biotechnology company, anchored in Japan, with a global reach

MISSION

Making a significant contribution to improving the quality of life and health of people around the world

VALUES

Integrity and Accountability, Passion, Courage and Resilience, Openness, Teamwork



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