

A Clear Investment Thesis

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Chief Executive Officer

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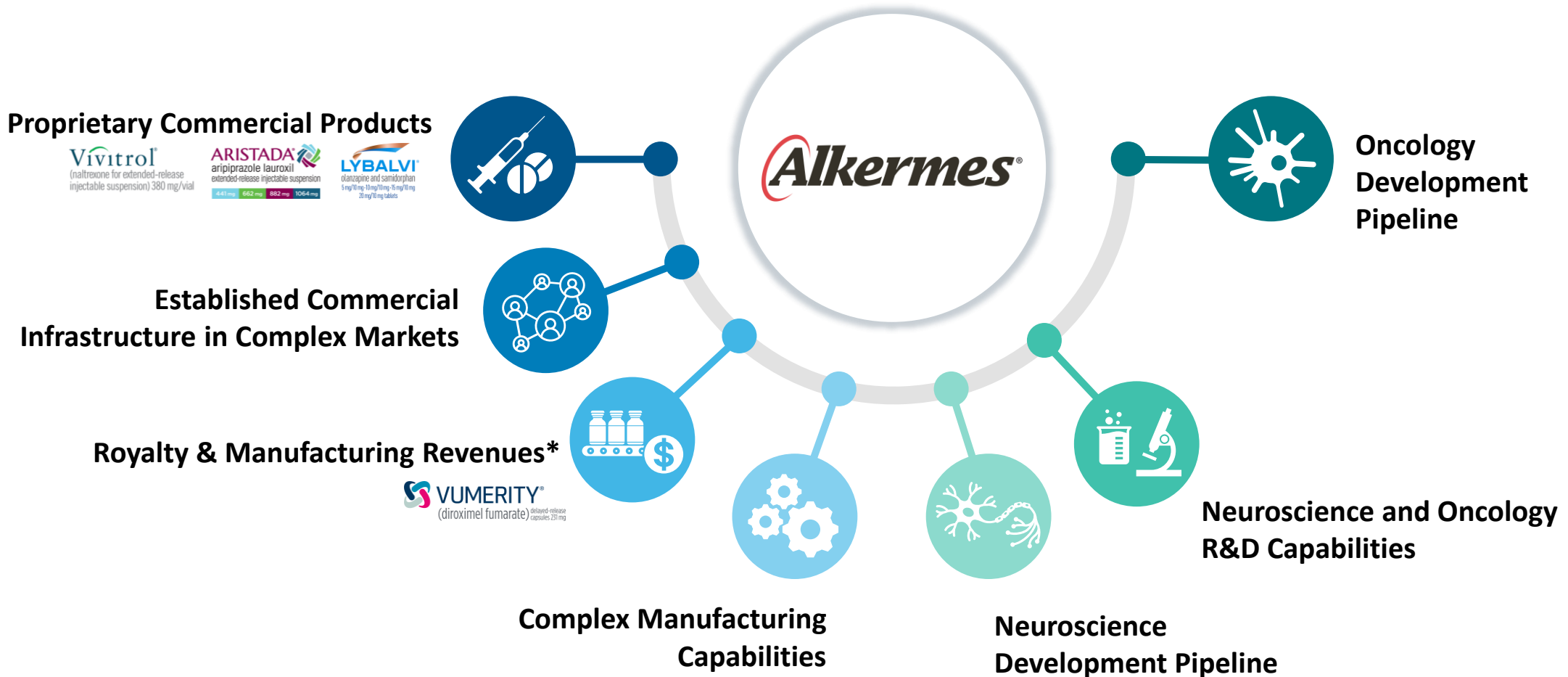


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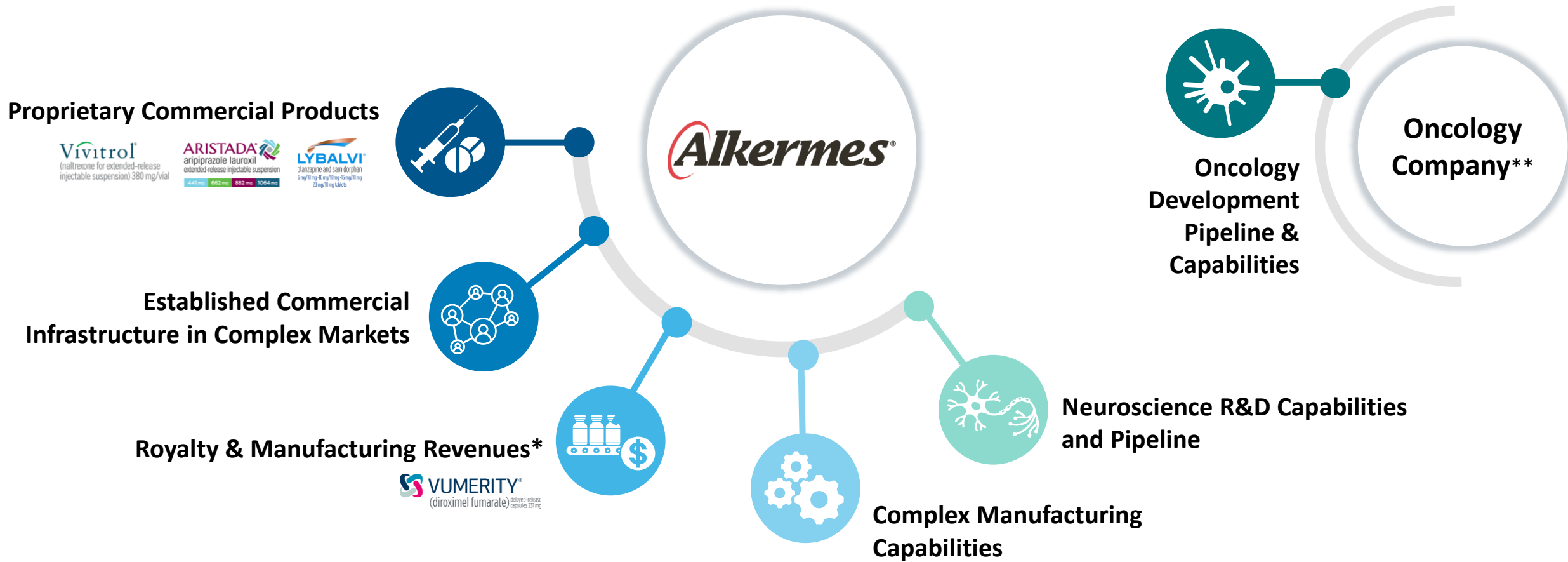
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Alkermes Today



*VUMERITY is licensed to and commercialized exclusively by Biogen

Post-Separation: Clear Value Propositions in Neuroscience and Oncology



*VUMERITY is licensed to and commercialized exclusively by Biogen

**Assuming separation of the company's oncology business is effected through a spin-off of the oncology business into an independent, publicly-traded company

Clear Priorities to Unlock Value in 2023

2023 Business Priorities and Value Drivers

1

Drive launch of LYBALVI®

Continued execution of commercial strategy and investment in DTC campaign

2

Advance orexin 2 receptor agonist

Establish initial safety and tolerability profile and generate initial clinical proof-of-concept data for ALKS 2680

3

Separate oncology business

Clarify value proposition for standalone neuroscience and oncology businesses

DTC: Direct-to-consumer

Drive Launch of LYBALVI®

LYBALVI® (olanzapine and samidorphan): Oral Treatment Option for Adults With Schizophrenia or Bipolar I Disorder

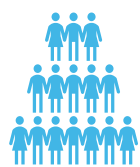


- Once-daily, oral atypical antipsychotic composed of olanzapine, an established antipsychotic agent, and samidorphan, a new chemical entity
- Commercially launched in U.S. Q4 2021 with differentiated label
- Indicated for the treatment of:
 - Schizophrenia in adults
 - Bipolar I disorder in adults
 - Acute treatment of manic or mixed episodes as monotherapy and as adjunct to lithium or valproate
 - Maintenance monotherapy treatment



Boxed Warning: Elderly patients with dementia-related psychosis treated with antipsychotic drugs are at an increased risk of death. LYBALVI is not approved for the treatment of patients with dementia-related psychosis. Full prescribing information may be found at www.lybalvi.com/lybalvi-prescribing-information.pdf

LYBALVI® Prescription Growth Trends Reflect Strong Prescriber Breadth and Diverse Source of Business



**>7,000
prescribers**

since launch (as of Nov. 30, 2022)

TRx indication split**:

~50% bipolar I disorder

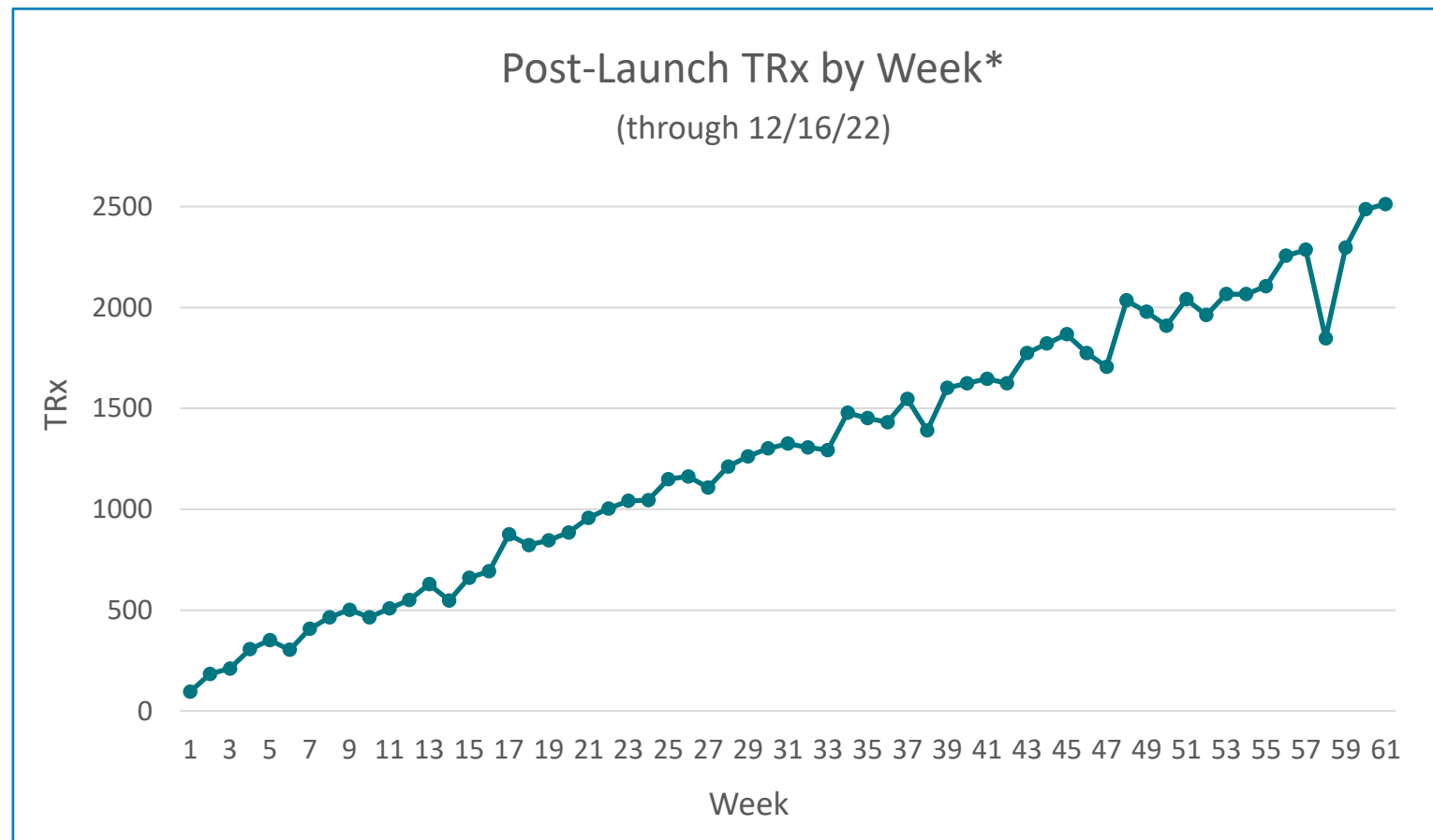
~50% schizophrenia

Patients switching from:

~45% olanzapine

~55% other branded and generic antipsychotic therapies

**Indication split based on claims data for patients with diagnoses of schizophrenia or bipolar I disorder only.



*Source: IQVIA

LYBALVI® Indications Represent a Large U.S. Opportunity

	Schizophrenia	Bipolar I Disorder†
Patients (Adults)	~2,600,000*	~3,900,000 - 5,200,000†
Monthly Treatment Switches	~22,000± 3,100 switches to branded oral agents	~31,000± 9,700 switches to branded oral agents
Payer Mix	82% Medicaid/Medicare±	64% Medicaid/Medicare±

*Desai et al. *J Manag Care Pharm.* 2013;19(6):468-77.

†Blanco et al. *J Psychiatr Res.* 2017 January; 84: 310–317.; Grant et al. *J Clin Psychiatry* 2005; 66: 1205-1215.

U.S. adults number based on 2020 U.S. Census Bureau population estimate (<https://www.census.gov/quickfacts/fact/table/US/PST045219> accessed on Jan 5, 2023)

±IQVIA Source of Business as of November 2022 R3M data

LYBALVI is indicated for the treatment of schizophrenia and bipolar I disorder in adults

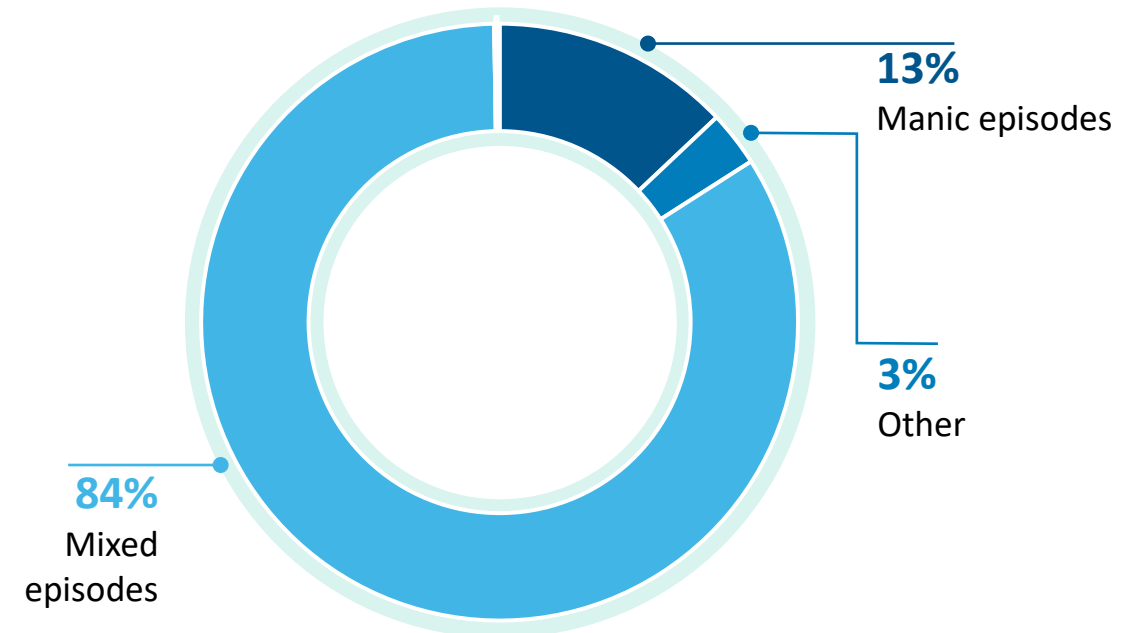
LYBALVI®: Bipolar I Disorder Label Relevant to Real-World Patients



	Acute treatment of manic episodes	Acute treatment of mixed episodes	Maintenance treatment
Adjunct to lithium or valproate	✓	✓	
Monotherapy	✓	✓	✓

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Healthcare providers (n=172) surveyed reported that majority of their bipolar I disorder patients faced mixed episodes (manic and depressive symptoms)



Survey question: What Bipolar I disorder symptoms is this patient facing, that you hope to treat with their current therapy regimen?

Source: Alkermes Market Survey; Fielded April 29 to June 2, 2022; N=172 HCPs (515 Patient Charts)

Plans to Launch LYBALVI® Direct-to-Consumer Advertising Campaign Focused on Bipolar I Disorder

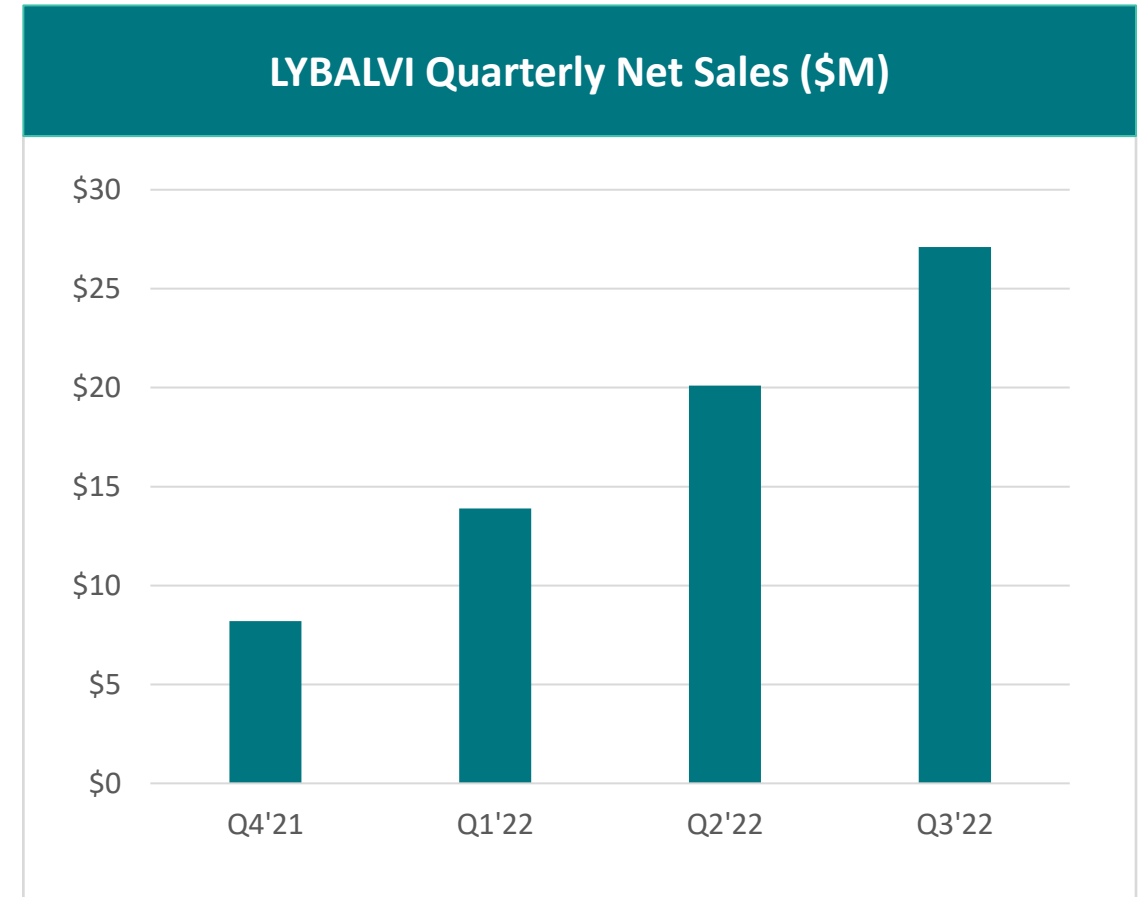


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LYBALVI®: Strong Launch Trajectory Established in 2022 With Clear Strategic Focus in 2023

2023 Commercial Strategy Focus

- **Breadth:** Continue to drive prescriber breadth through highly-targeted field force deployment
- **Access:** Expand patient access through continued execution of disciplined payer contracting strategy
- **Awareness:** Launch DTC advertising campaign focused on digital and broadcast channels

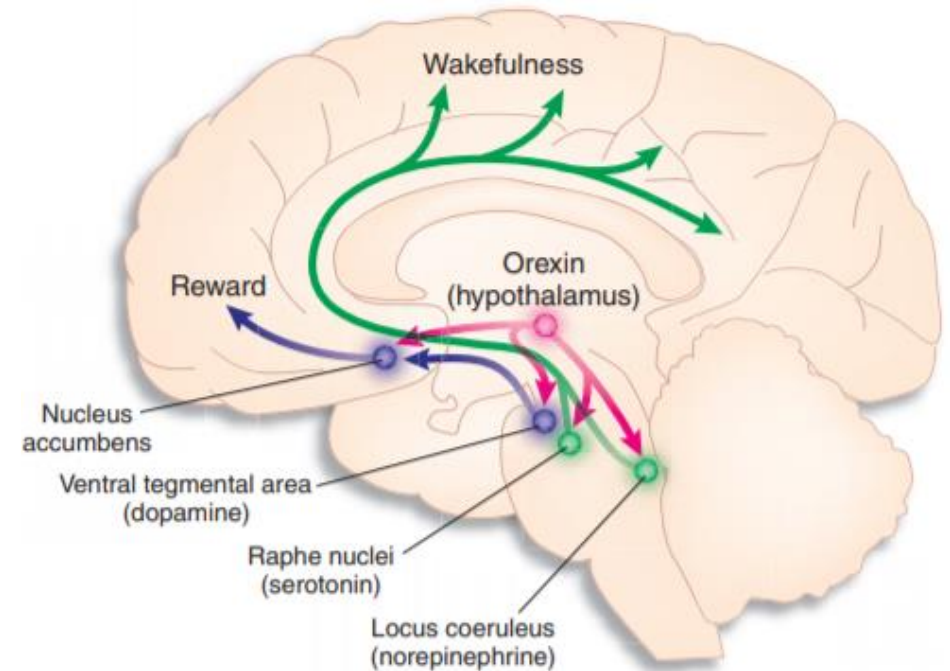




Advance Orexin 2 Receptor (OX2R) Agonist

Orexin Dysfunction: Well Defined Opportunity in Narcolepsy and Other Sleep Disorders

- In narcolepsy and other sleep disorders, low orexin levels lead to inconsistent neurotransmitter release, resulting in excessive sleepiness and poor regulation of REM sleep
- Narcolepsy affects ~200,000 people in U.S. and 3M people globally¹
- 70% of people with narcolepsy have narcolepsy type 1², distinguished by:
 - Cataplexy, a sudden muscle weakness triggered by strong emotions
 - Low or no orexin in the brain
- Genetic and pharmacologic evidence suggests that orexin receptor agonists, especially OX2R agonists, may be useful for mechanistic therapy of narcolepsy³



¹ Global Narcolepsy Drugs Market, Forecast 2019-2025. Allied Market Research

² Swick TJ. Treatment paradigms for cataplexy in narcolepsy: past, present, and future. *Nat Sci Sleep*. 2015;7:159-169

³ Nagahara T. Design and Synthesis of Non-Peptide, Selective Orexin Receptor 2 Agonists. *J. Med. Chem.* 2015;58:7931-7937

Figure from: Scammell, T E, and Saper, C B. *Nature medicine*. 2007;13:126-8

Leveraging Alkermes' Molecular Design Capabilities to Target Orexin Dysfunction

ALKS 2680 molecular design objectives:

- Capture performance of endogenous peptide OX2R agonist
 - Increased wakefulness duration
 - Improved cataplexy control
- Optimize potency and minimize predicted human dose to mitigate risk of undesired side effects
- Provide PK/PD profile that mirrors natural wake cycle
 - Dose to allow for 8-12 hours wakefulness without subsequent insomnia
- Enable convenient once-daily, oral medication

Orexin 2 Receptor in Complex with Peptide-Agonist Orexin-B

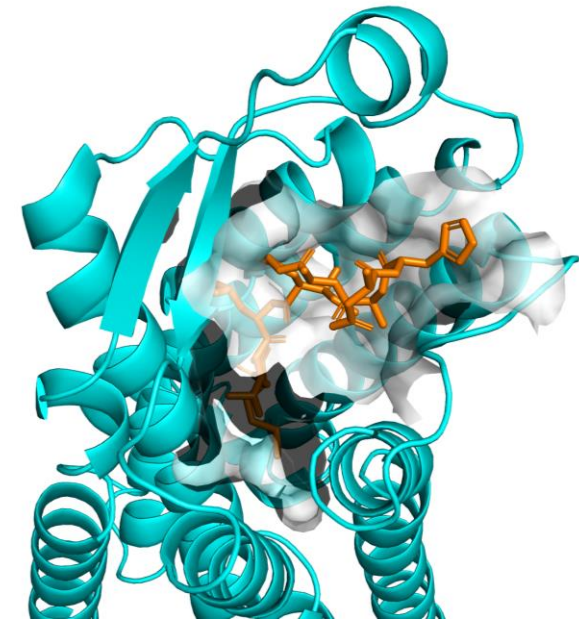
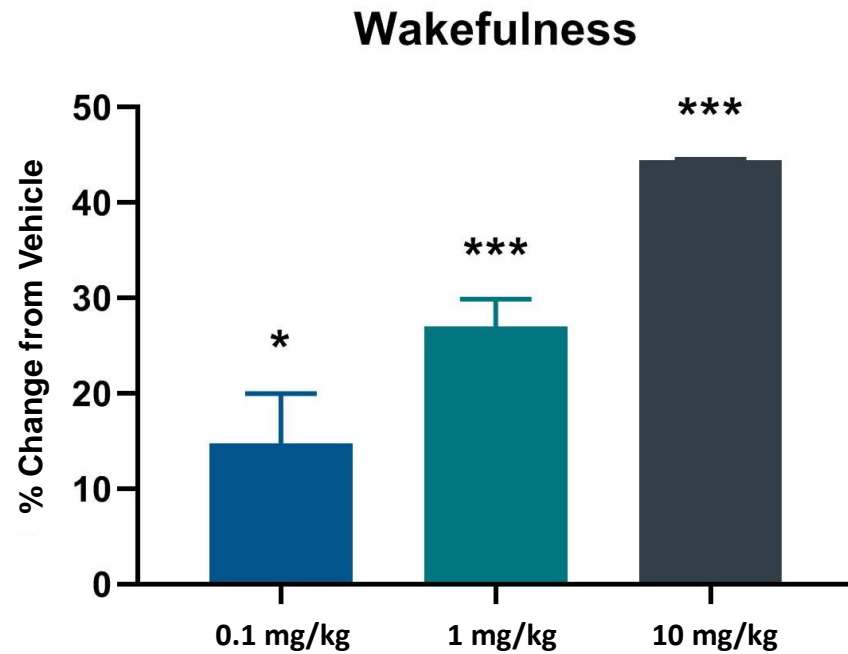


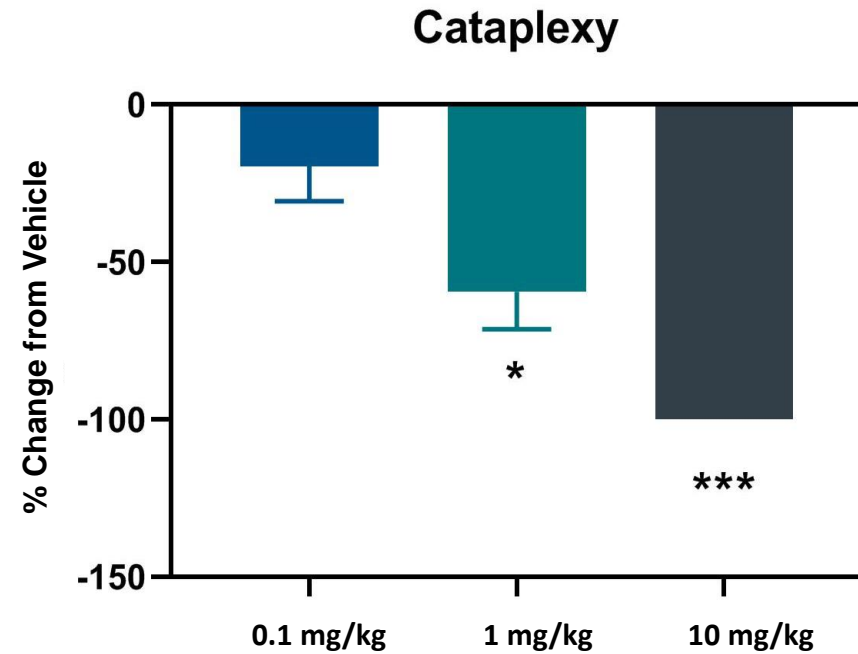
Figure adapted from: Hong, Chuan, et al. *Nature communications*. 2021:12; 3. PDB ID: 7L1U

Selective OX2R Agonist Showed Dose-Dependent Increased Wakefulness and Reduced Cataplexy in Preclinical Models



RDC-264177

Mean ± SEM; One-way ANOVA
* = p<0.05, *** = p<0.001



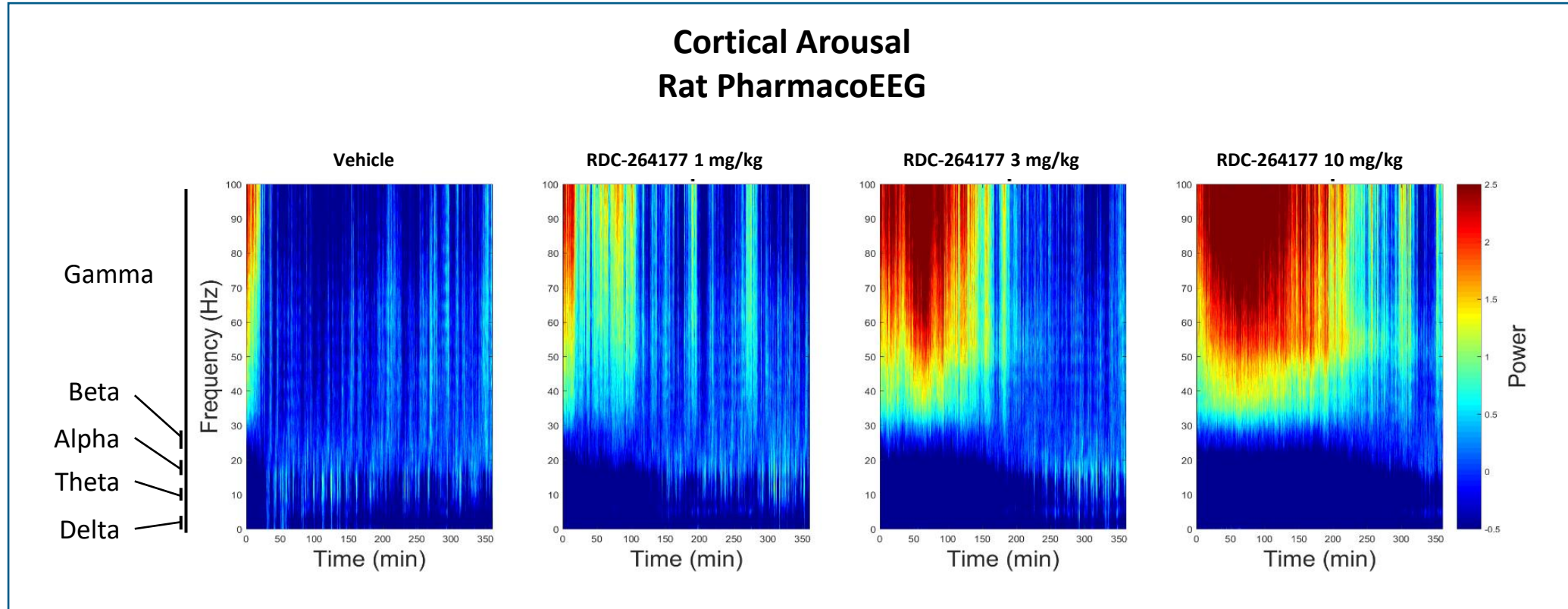
RDC-264177

DTA mouse model of narcolepsy^{1,2} believed to serve as a predictive disease model of narcolepsy in humans

¹Tabuchi S, Tsunematsu T, Black SW, et al. Conditional ablation of orexin/hypocretin neurons: a new mouse model for the study of narcolepsy and orexin system function. *J Neurosci*. 2014;34(19):6495-6509

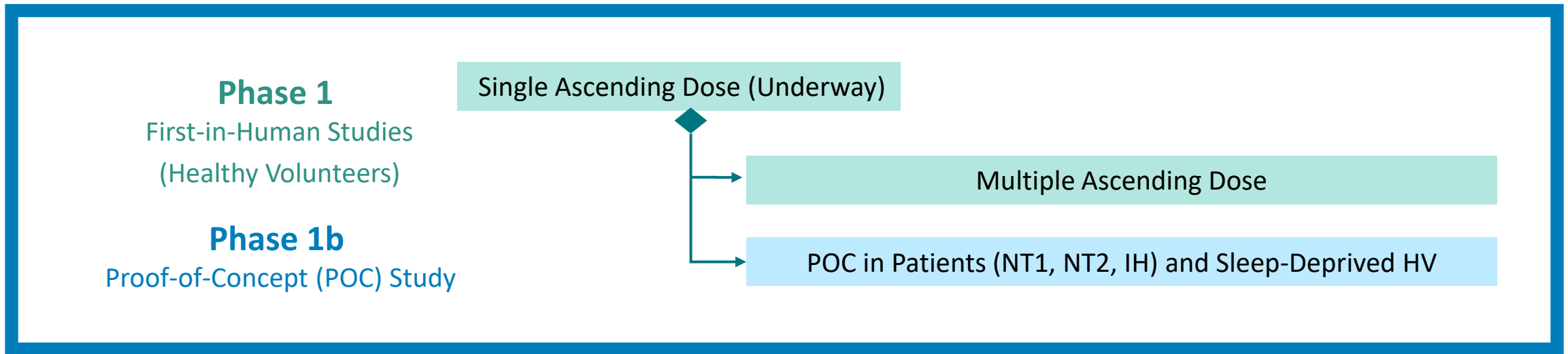
² In collaboration with SRI International

Selective Orexin 2 Receptor Agonist Promoted Prolonged Wakefulness in Preclinical *In Vivo* Studies



Selective orexin 2 receptor agonist demonstrated dose-dependent wake duration
(shown by red gamma bands and dark blue theta and delta bands)

ALKS 2680: Clinical Development Plan Designed to Rapidly Advance Program in 2023



NT1: Narcolepsy Type 1; **NT2:** Narcolepsy Type 2; **IH:** Idiopathic Hypersomnia; **HV:** Healthy Volunteers

ALKS 2680 Clinical Development Plan

Targeting Clinical Proof-of-Concept Data by Year-End 2023

Phase 1 First-in-Human (FIH) Studies

- Objective: Measure and model pharmacokinetics (PK)/ pharmacodynamics (PD) and evaluate safety and tolerability of single and multiple ascending doses
- Key Assessments:
 - Drug exposure (PK)
 - Evaluate safety and tolerability of single and multiple ascending doses
- Exploratory assessment of target engagement: qEEG trends in power of frequency bands
- Single-ascending dose study ongoing; Multiple-ascending dose study initiation expected Q1 2023

Phase 1b Proof-of-Concept (POC) Study

- Objective: Early POC data + dose range estimation for phase 2 in lead indications
- Key Assessments:
 - EEG-based maintenance of wakefulness test as primary efficacy/PD readout
 - Drug exposure (PK)
 - Evaluate safety and tolerability
- Study initiation expected H1 2023; Preliminary data expected by year-end

EEG: electroencephalogram; qEEG: quantitative electroencephalogram

Separate Oncology Business

Post-Separation Oncology Co.

Pure-Play, Development-Stage Oncology Company

Investment thesis anchored by potential medical and economic value of nemvaleukin alfa:

- Potential first-in-class IL-2 variant immunotherapy
- Anti-tumor activity observed both as a single agent and with checkpoint inhibitors (CPI), in CPI-unapproved tumor types and post-CPI settings
- Potential registration-enabling studies underway in mucosal melanoma* and platinum-resistant ovarian cancer**, each with FDA Fast Track Designation
- Investigating alternative routes of administration/dosing schedules

Sophisticated protein engineering platform capabilities and early-stage development assets

- Tumor-targeted split IL-12 program
- IL-18 program

Nemvaleukin **offers an opportunity for significant value creation** as the development program advances and expands

Highly-experienced team with **scientific and clinical trial expertise** to efficiently advance pipeline

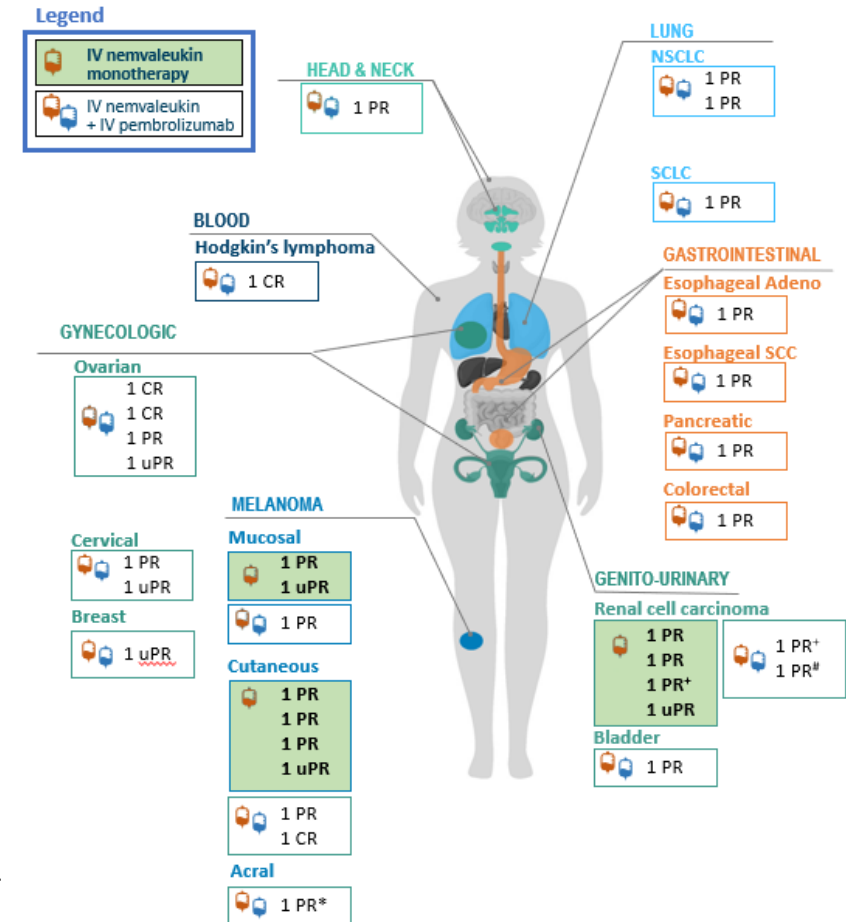
Opportunity to attract **oncology-focused investors**

*Also granted FDA Orphan Drug Designation; **In combination with pembrolizumab
Assuming separation is effected through a spin-off of the oncology business into an independent, publicly-traded company

ARTISTRY-1 (IV Nemvaleukin): Durable Responses Observed

- Demonstrated **durable responses** in high unmet need populations
 - **Monotherapy activity** (IV) in prior anti-PD-(L)1 treated melanoma and renal cell carcinoma
 - **Combination activity** (IV) with pembrolizumab in a range of tumor types
- Treatment-related adverse events (AEs) have been consistent with expectations based on nemvaleukin's mechanism of action and were mostly transient and manageable
 - Pyrexia, chills and nausea were the most commonly reported AEs; Transient and asymptomatic neutropenia/neutrophil count decrease were the most commonly reported events of grade ≥ 3
 - Three dose-limiting toxicities were reported, all in the highest dose evaluated (declared as the maximum tolerated dose)

Monotherapy and Combination Responses



Patients achieved SD (*acral), PR (+RCC), and PD (#RCC) on nemvaleukin monotherapy, rolled over to combination therapy and achieved PR. IV: Intravenous; NSCLC: Non-small cell lung cancer; SCLC: Small cell lung cancer; RCC: Renal cell carcinoma; SCC: Squamous cell carcinoma CR: complete response; PR: partial response; uPR: unconfirmed PR

Data as of August 2022

Nemvaleukin ARTISTRY Development Program

Potential Registration Enabling Studies

ARTISTRY-6

Advanced cutaneous and mucosal melanoma

- Phase 2
- Dosing: Daily IVx5, SC Q1W
- Monotherapy

Status

- IDMC: Risk/benefit profile supports study continuation as planned
- Mucosal melanoma enrollment ongoing
- Cutaneous cohort (SC) enrollment nearing completion
- Plan to initiate POC cohort with LFIV dosing (cutaneous)

ARTISTRY-7

Platinum-resistant ovarian cancer

- Phase 3
- Dosing: Daily IVx5
- Combination with pembrolizumab

Status

- Site initiation and enrollment ongoing
- In collaboration with MSD
- In partnership with the GOG Foundation and ENGOT

Alternative Dosing Studies

ARTISTRY-2

Advanced solid tumors

- Phase 1/2
- Dosing: SC Q1W
- Combination with pembrolizumab

Status

- Enrollment closed
- Data maturing, focus on durability of responses compared to daily IVx5
- Data from ARTISTRY-2 and ARTISTRY-6 will inform viability of SC

ARTISTRY-3

Advanced solid tumors

- Phase 1/2
- Dosing: LFIV dose escalation
- Monotherapy and combination with pembrolizumab

Status

- Enrolling Day 1 Q3W and Day 1,4 Q3W dosing cohorts at pharmacologically relevant doses
- Escalation ongoing

IV: Intravenous; LFIV: Less frequent IV; SC: Subcutaneous; IDMC: Independent Data Monitoring Committee; MSD: A tradename of Merck & Co., Inc. Kenilworth, NJ, USA; ENGOT: European Network of Gynaecological Oncological Trial Groups



Positioning Alkermes Neuroscience Business for Future Growth

Post-Separation Alkermes*

Pure-Play, Commercial-Stage Neuroscience Company

Builds on Alkermes' innovation and excellence in neuroscience



Proprietary Products

- Topline primarily driven by growth of proprietary commercial products in addiction and psychiatry

Vivitrol
(naltrexone for extended-release injectable suspension)

ARISTADA
aripiprazole lauroxil
extended-release injectable suspension

LYBALVI
olanzapine and samidorphan

- Complex manufacturing capabilities



Commercial Capabilities

- Established commercial capabilities in complex psychiatry and addiction markets
- Opportunity to capture further operating leverage



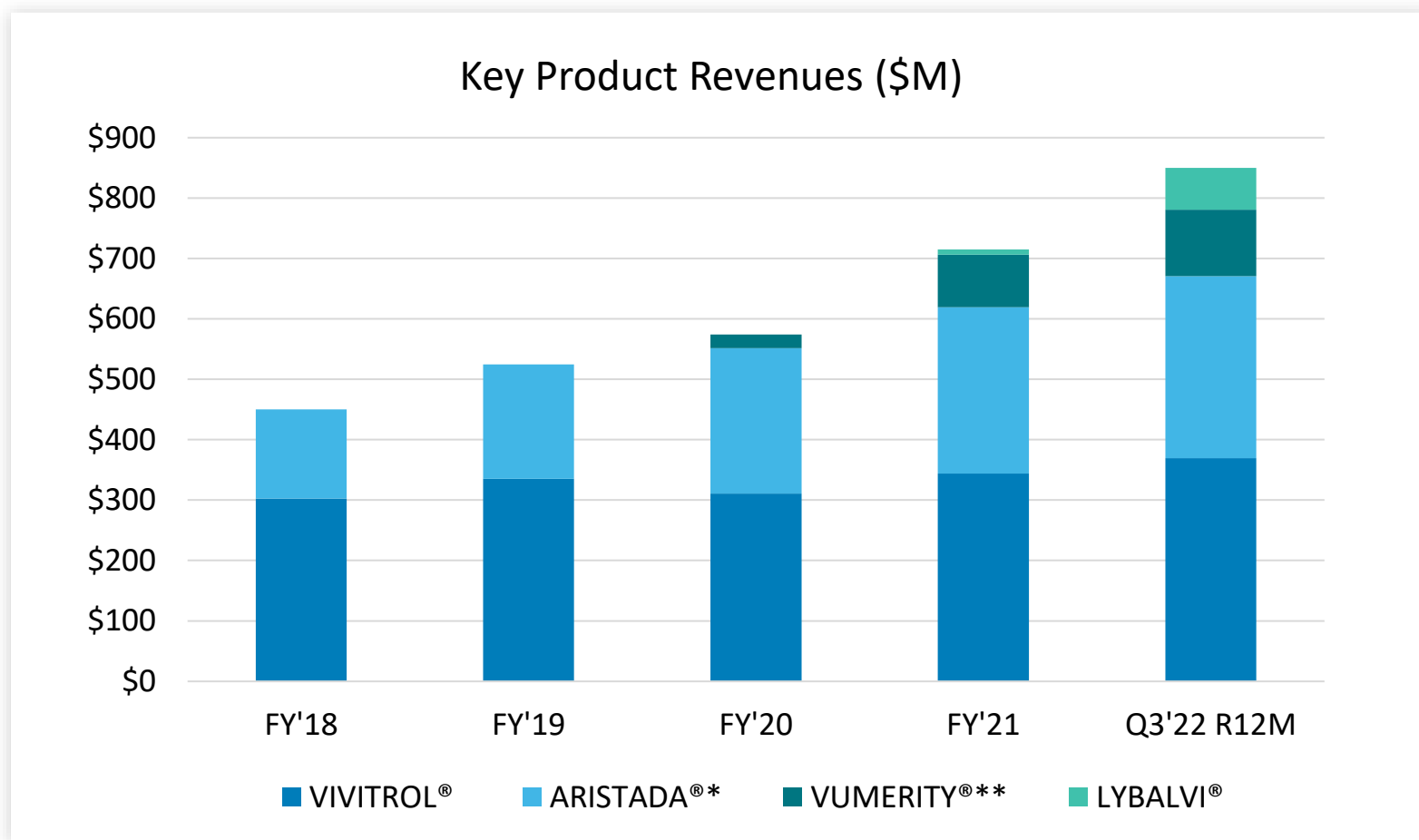
Development Pipeline

- Early-stage neuroscience pipeline
 - ALKS 2680, orexin 2 receptor agonist in phase 1
 - Portfolio of preclinical neuroscience assets

Separation expected to enhance profitability

*Assuming separation of the company's oncology business is effected through a spin-off of the oncology business into an independent, publicly-traded company

Topline Growth and Diversification Reflect Evolving Business



- Plan to provide 2023 financial expectations and updated long-term profitability targets in Q1'23

*Inclusive of ARISTADA INITIO®

**Licensed product (royalty & manufacturing revenue)

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Establish initial safety and tolerability profile and generate initial clinical proof-of-concept data for ALKS 2680

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Separate oncology business

Clarify value proposition for standalone neuroscience and oncology businesses

www.alkermes.com

