



# Corporate Presentation

A Breakthrough Neuropsychiatry Company

August 2025

NYSE American: CYBN

Cboe CA: CYBN

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## DRUG DEVELOPMENT

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# Changing Minds: Redefining the Standard of Care in Mental Health

Cybin is developing **differentiated, next-generation therapeutics** with the potential to **improve clinical outcomes and address key unmet needs** for people with mental health conditions<sup>1</sup>

## A Novel Treatment Approach to Neuropsychiatry

Cybin is advancing **intermittent treatments** with potential **rapid-onset, long-lasting clinical efficacy** in treating depression and anxiety

Unlike current treatments that only address symptoms, our therapies **target underlying causes in neural circuitry** that lead to mental health disorders

# We Are an Advanced Clinical-Stage Neuropsychiatry Company Approaching Key Near-Term Milestones<sup>1</sup>

1

**Two proprietary clinical programs, CYB003 and CYB004,** targeting depression and anxiety disorders with **positive Phase 2 safety and efficacy results**

2

Lead program CYB003, which has been granted **U.S. Food and Drug Administration Breakthrough Therapy Designation** is in **Phase 3 development for the adjunctive treatment of Major Depressive Disorder (“MDD”)**

3

**Robust pipeline of differentiated assets** with potential for expansion into **additional neuropsychiatry indications with high unmet need affecting >200M people in the U.S.**<sup>2</sup>

4

**Strong Intellectual Property Portfolio:** over 95 granted patents, over 250 patent applications pending

5

**Well-Capitalized** to move programs forward with cash position of **US\$118.7 million** as of June 30, 2025

# Exceptional Team Pedigree With Successful Track Record of Bringing Drugs to Market<sup>1</sup>



**Doug Drysdale**  
Chief Executive  
Officer



**Amir Inamdar MBBS,  
DNB(Psych),FFPM**  
Chief Medical Officer



**Alex  
Nivorozhkin, Ph.D**  
Chief Scientific Officer



**Aaron Bartlone**  
Chief Operating  
Officer



**Tom Macek**  
SVP, Clinical  
Development



**Atul R.  
Mahableshwarkar, M.D.,  
DLFAPA**  
SVP, Clinical  
Development



**Allison  
House-Gecewicz**  
SVP, Clinical  
Operations



**Geoff Varty  
Ph.D.**  
Head of  
R&D

- Combined 60 Investigational New Drug (IND) applications, 37 exits
- Combined 300 peer-reviewed publications by scientific leadership

**Trintellix**

**Celexa**  
citalopram HBr

**Lexapro**  
escitalopram oxalate

**Namenda**  
memantine HCl tablets

**Sabril**

**Allegra**

**Anzemet**  
dolasetron mesylate injection/tablets

**Latuda**  
(lurasidone HCl)

**Rozerem**  
ramelteon 8-mg tablets

**VANIQA**

**zolgensma**

**Saphris**

# Innovative Neuropsychiatry Pipeline of 5-hydroxytryptamine ("5-HT") Receptor Agonists with Clinical Validation and Value-Driving Milestones

PROGRAM	INDICATION	DISCOVERY	PRE CLINICAL	PHASE 1	PHASE 2	PHASE 3	NEXT MILESTONES <sup>1,2,3</sup>
<b>CYB003</b> Deuterated Psilocin (Oral)	Adjunctive treatment of Major Depressive Disorder (MDD)	Phase 3 study dosing underway Granted FDA Breakthrough Therapy Designation					<b>Mid-2025:</b> initiate second pivotal study, EMBRACE
<b>CYB004</b> Deuterated Dimethyltryptamine (Intramuscular)	Generalized Anxiety Disorder (GAD)	Phase 2 study dosing underway					<b>August 2025:</b> Phase 2 GAD study expected to complete enrollment
<b>CYB005</b> Phenethylamines (Non-hallucinogenic doses)	CNS Disorders	Preclinical studies underway					



Notes:

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3) Cybin is prioritizing the progression of its CYB003 program. The advancement of Cybin's CYB004, CYB005 and technology programs are all contingent on Cybin's ability to continue raising capital under its current and future financing arrangements. No assurances can be given that Cybin will be able to raise the additional capital that it may require for its anticipated future development.

# Pipeline Targets Neuropsychiatry Indications with High Unmet Need

	Addressable Market	Health Impact	Need for Improved Treatments
<u>CYB003</u> MDD	<p>&gt; 300 million people worldwide<sup>1</sup></p> <p>~21 million with MDD in the U.S.<sup>2</sup></p>	<ul style="list-style-type: none"> <li>• Suicide risk is 20x higher for an individual with vs. without depression<sup>3</sup></li> <li>• 50-75% of MDD patients have anxious depression<sup>4</sup></li> </ul>	<ul style="list-style-type: none"> <li>• 2/3rds of patients do not experience relief with initial antidepressant treatment<sup>5</sup></li> <li>• SSRI/SNRI side effects: weight gain (20%)<sup>6</sup>, sexual dysfunction (up to 30%)<sup>7</sup>, GI disturbances* and insomnia (25%)<sup>8</sup></li> <li>• With 2<sup>nd</sup> and 3<sup>rd</sup> line treatments, efficacy decreases; intolerance and relapse rates increase<sup>9</sup></li> </ul>
<u>CYB004</u> GAD	<p>&gt; 300 million people with anxiety disorders worldwide<sup>10</sup></p> <p>6.8 million with GAD in the U.S. (3.1% of population)<sup>11</sup></p>	<ul style="list-style-type: none"> <li>• GAD is the most common anxiety disorder seen in primary care<sup>12</sup></li> <li>• ~77% of adults with GAD have moderate to severe impairment<sup>13</sup></li> </ul>	<ul style="list-style-type: none"> <li>• 50% of patients with GAD do not respond to first line treatment with SSRIs and SNRIs<sup>12</sup></li> <li>• 57% of patients with anxiety do not adhere to SSRI/SNRIs, due to side effects<sup>14</sup></li> </ul>

# Transforming the Treatment Paradigm for Mental Health<sup>5,6</sup>

## Barriers to accessing care with current treatments:

### Low availability to see new patients

- **Only 18.5%** of U.S. psychiatrists available to see new patients<sup>1</sup>

### Long median wait times<sup>1</sup>

- 67 days for in-person visits, 43 days for telepsychiatry

### High frequency of visits for existing intermittent treatments

- Esketamine: 26 sessions<sup>2</sup>
- TMS: Total of up to 36 sessions (5 per week)<sup>3</sup>
- ECT: Total of 6-12 sessions (2-3 per week)<sup>4</sup>



**CYB003** with **infrequent acute dosing** and **long-lasting relief** presents opportunity to:

- **Reduce** frequency of visits for existing patients
- **Lower barriers to timely care**

#### Notes:

1) Sun et al. (2023). Low availability, long wait times, and high geographic disparity of psychiatric outpatient care in the US. *General Hospital Psychiatry*, 84, 12–17. <https://doi.org/10.1016/j.genhosppsych.2023.05.012>

2) Esketamine package insert

3) Hutton et al. (2023). Dosing transcranial magnetic stimulation in major depressive disorder: Relations between number of treatment sessions and effectiveness in a large patient registry. *Brain stimulation*, 16(5), 1510–1521. <https://doi.org/10.1016/j.brs.2023.10.001>

4) Thirthalli, J., Naik, S. S., & Kunigiri, G. (2020). Frequency and Duration of Course of ECT Sessions: An Appraisal of Recent Evidence. *Indian journal of psychological medicine*, 42(3), 207–218. [https://doi.org/10.4103/IJPSYM.IJPSYM\\_410\\_19](https://doi.org/10.4103/IJPSYM.IJPSYM_410_19)

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# Opportunity for Pipeline Expansion into Indications with Large Addressable Markets<sup>15,16</sup>

	Indications with early studies supporting potential	U.S. Prevalence	Estimated Addressable Market*
Behavioral Disorders	Depression <sup>1</sup>	8.3% <sup>8</sup>	28 million
	Anxiety Disorders / PTSD <sup>2</sup>	19.1% / 3.6% <sup>9</sup>	64 million / 12 million
	Bipolar Disorder <sup>3</sup>	2.8% <sup>10</sup>	9 million
	Substance Use / Addiction Disorders <sup>4</sup>	14.5% <sup>11</sup>	48 million
	Eating Disorders <sup>5</sup>	0.3-1.2% <sup>12</sup>	1-4 million
Other CNS	Cluster Headaches / Migraine <sup>6</sup>	0.1% / 12% <sup>13</sup>	0.3 million / 40 million
	Chronic Pain Management <sup>7</sup>	21% <sup>14</sup>	70 million



Large opportunity for expansion into indications **affecting >200 million people in the U.S.**

# CYB003

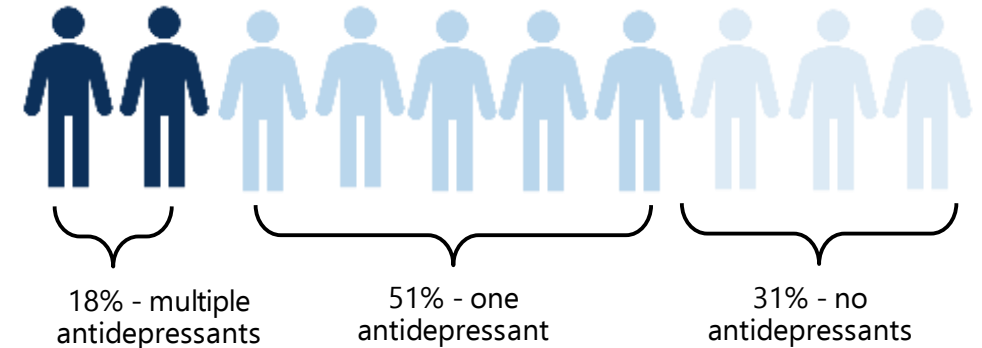
Deuterated Psilocin Program  
Adjunctive Treatment of MDD

# CYB003: An Acute Adjunctive Therapy with Potential for Durable Remission

## An adjunctive therapy could potentially<sup>1</sup>:

- ✓ Allow for immediate treatment without waiting to withdraw from background medications
- ✓ Prevent withdrawal symptoms, which could be severe for some patients after years of antidepressant use
- ✓ Eliminate logistical hurdles associated with titrating off existing medications
- ✓ Allow patients to retain some benefit from background medications even if the background medications are inadequate alone

## Current MDD Treatment in the U.S.<sup>2</sup>



Majority of patients are being treated with background medication<sup>2</sup>

~70% of patients on SSRIs<sup>3</sup>/SNRIs<sup>3</sup>  
~60% on antidepressant > 2 years  
~44% on antidepressant > 5 years

# CYB003 Program Overview

## Novel treatment paradigm

- Intermittent dosing with rapid relief and long-lasting remission
- Adjunctive treatment for patients who do not experience relief with SS RIs

## Best-in-class effect size

- Primary endpoint at 3 weeks: -13.75 point difference in change in Montgomery-Asberg Depression Rating Scale ("MADRS") from baseline between CYB003 (12mg and 16mg pooled) vs. placebo ( $p < 0.0001$ )

## Robust, sustained efficacy at 12 months

- **100%** of participants were responsive to treatment and **71%** were in remission from depression 12 months after 2 doses (16 mg) in a Phase 2 study
- Mean ~23-point reduction in MADRS scores from baseline

## Breakthrough Therapy Designation ("BTD")

- U.S. FDA BTD for adjunctive treatment of MDD
- Phase 2 data for CYB003 shows preliminary evidence of significant improvements over existing therapies

## Upcoming Milestones<sup>1,2,3</sup>

- Initiate second pivotal Phase 3 study (EMBRACE) around mid-2025

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# Positive Phase 2 CYB003 Results in MDD

Rapid onset of effect

Improvement in symptoms after single dose

Large improvements in symptoms

At 3 weeks: 12 mg better than placebo on MADRS by 14.1 points ( $p=0.0001$ ), Cohen's  $d=2.31$   
16 mg better than placebo on MADRS by 13 points ( $p=0.008$ ), Cohen's  $d=2.54$

Incremental benefit of 2<sup>nd</sup> dose

Average 5.8 points improvement on the MADRS after 2<sup>nd</sup> dose (12 mg)  
>75% response rates and up to 79% remission rates (12 mg) after a 2<sup>nd</sup> dose

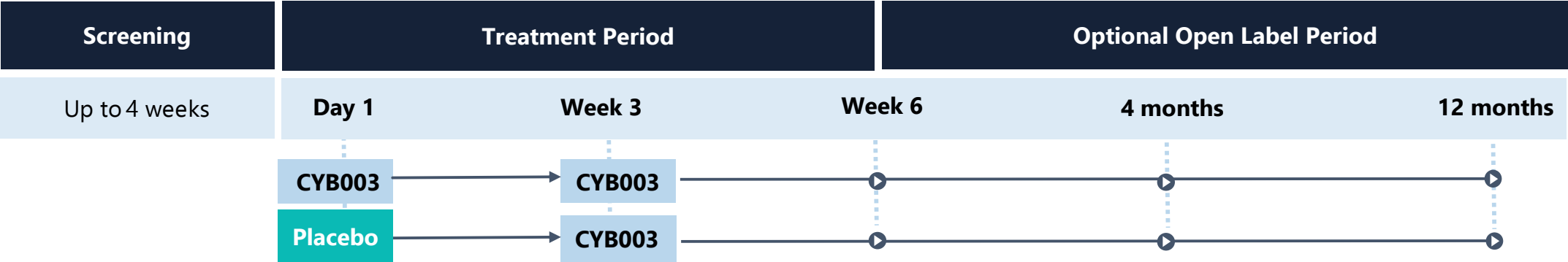
Durable efficacy at 12 months

Benefit sustained to 12 months with 71% remission rate and 100% response rate after 2 doses (16 mg)

Favorable safety and tolerability profile

All reported AEs<sup>1</sup> mild to moderate; no AEs of suicidality. No AEs reported at 12 months.

# CYB003: Phase 2a Trial Design in MDD <sup>1,2,3</sup>



**Phase 1:** Single ascending dose study (1-10 mg), n=12

**Phase 2a:** RCT in MDD patients (12 mg, n=24; 16 mg, n=12)

**Key Inclusion Criteria:**

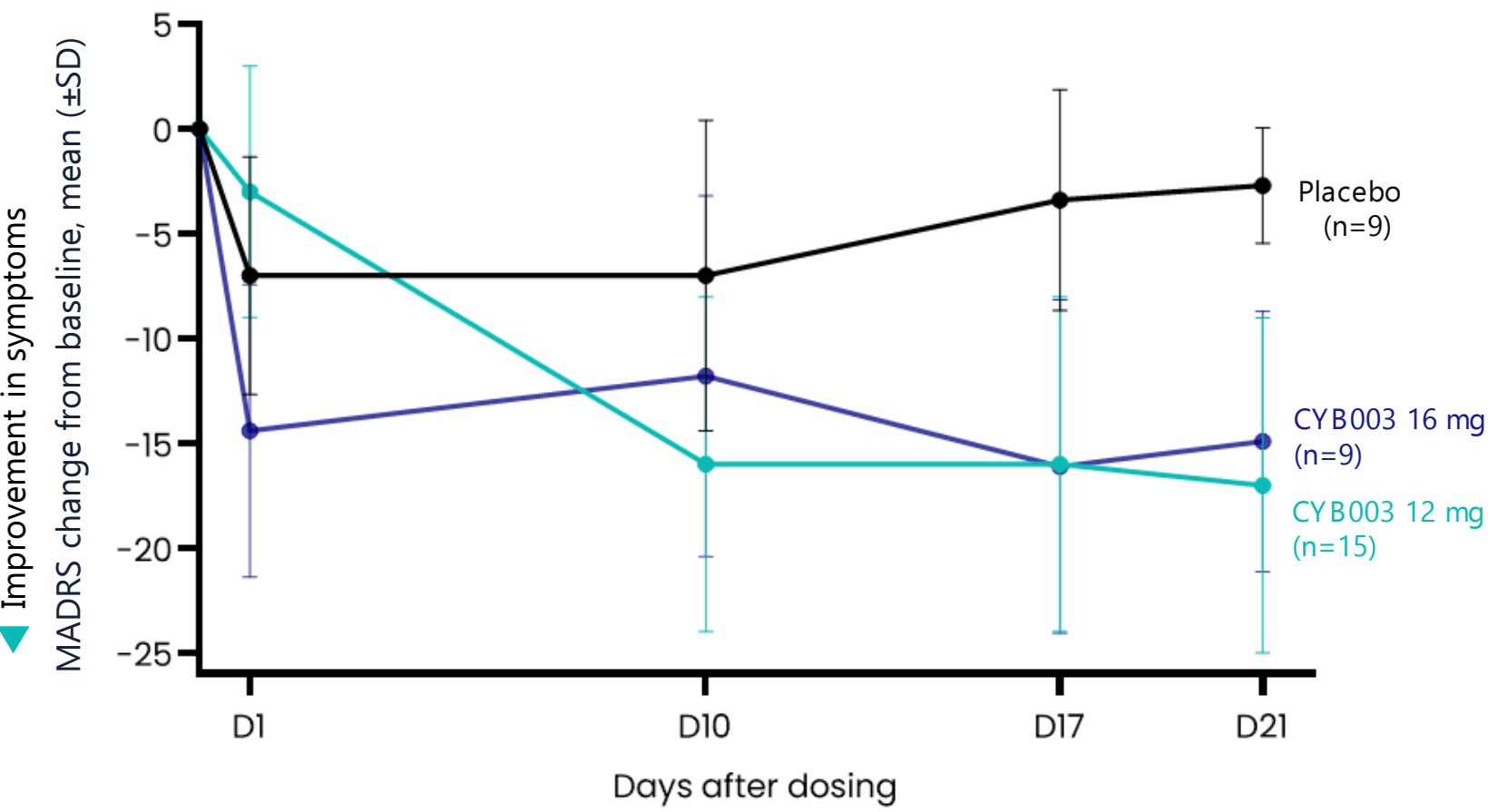
- ✓ Moderate to severe MDD (MADRS ≥ 21)
- ✓ Inadequate response to antidepressant medication

**Primary Endpoint:**

- ✓ Reduction in depression symptoms (change in MADRS score) at Week 3 after a single dose\* vs. placebo

Notes:  
1) \*Patients allowed to remain on stable doses of antidepressant medications  
2) Primary efficacy assessed at Week 3; Optional 12 week follow up to assess durability of effects  
3) Forward looking statements are subject to risks and assumptions. See “Cautionary Statement” on page 2 of this presentation. There is no assurance that timelines will be met. Anticipated timelines regarding drug development are based on reasonable assumptions informed by current knowledge and information available to Cybin. Such statements are informed by, among other things, regulatory guidelines for developing a drug with timeline safety studies, proof of concept studies, and pivotal studies for new drug application submission and approval, and assumes the success of implementation and results of such studies on timelines indicated as possible by such guidelines, other industry examples, and Cybin’s development efforts to date.

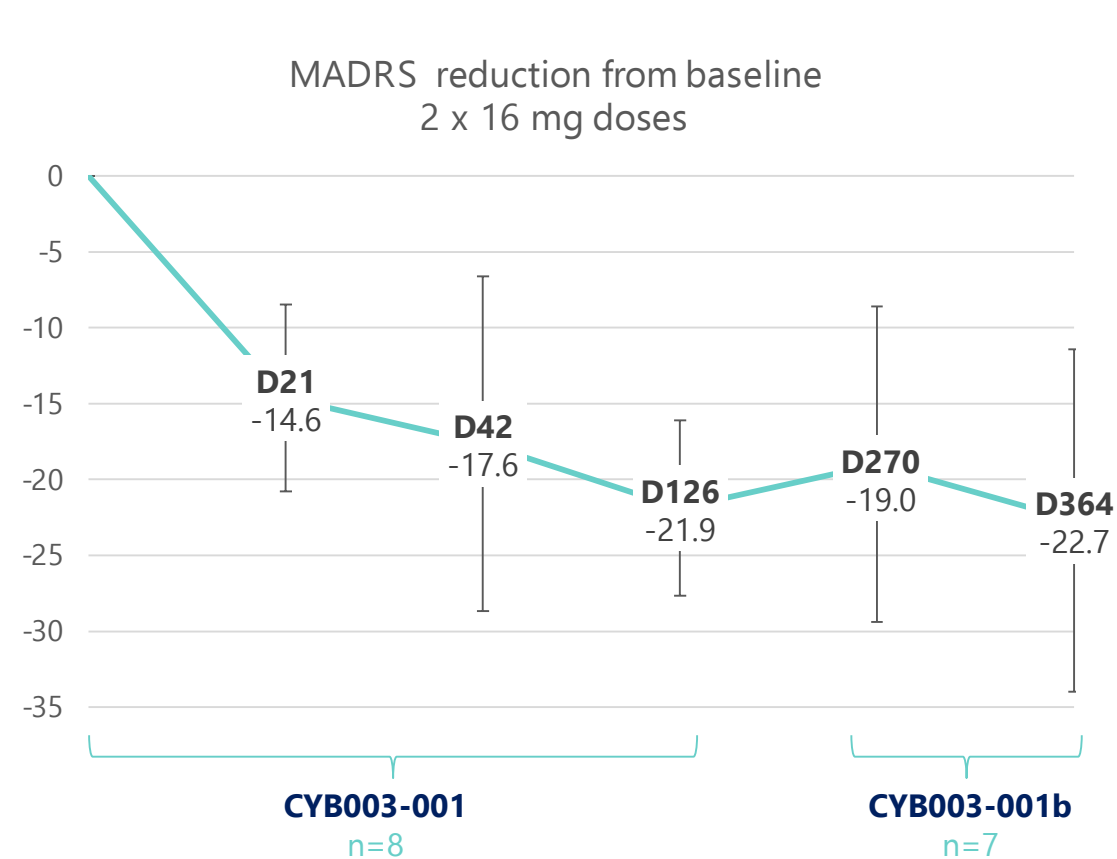
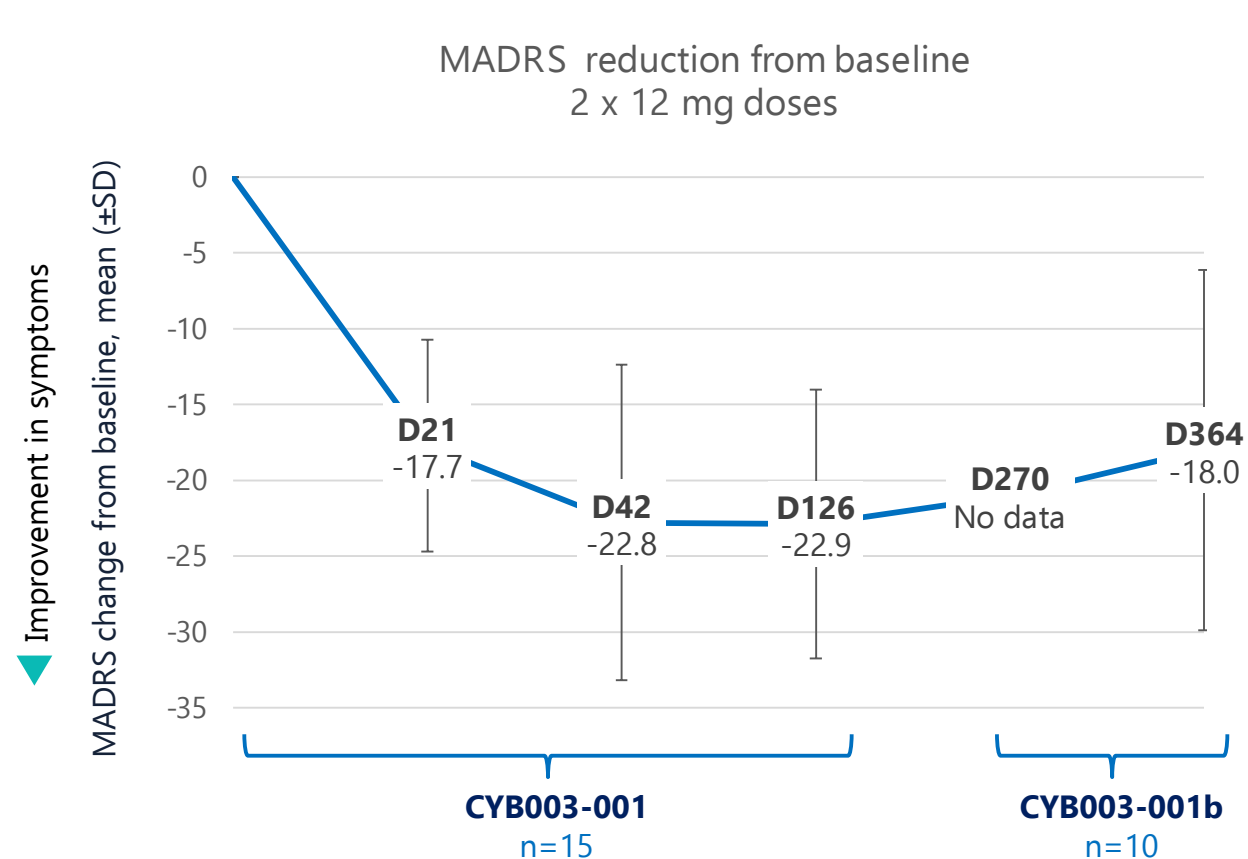
# Large Improvement in Depression Symptoms After a Single Dose of CYB003



Dose	Primary Endpoint*	Effect size	p-value
12 mg	-14.1	2.31	0.0001
16 mg	-12.99	2.54	0.0080

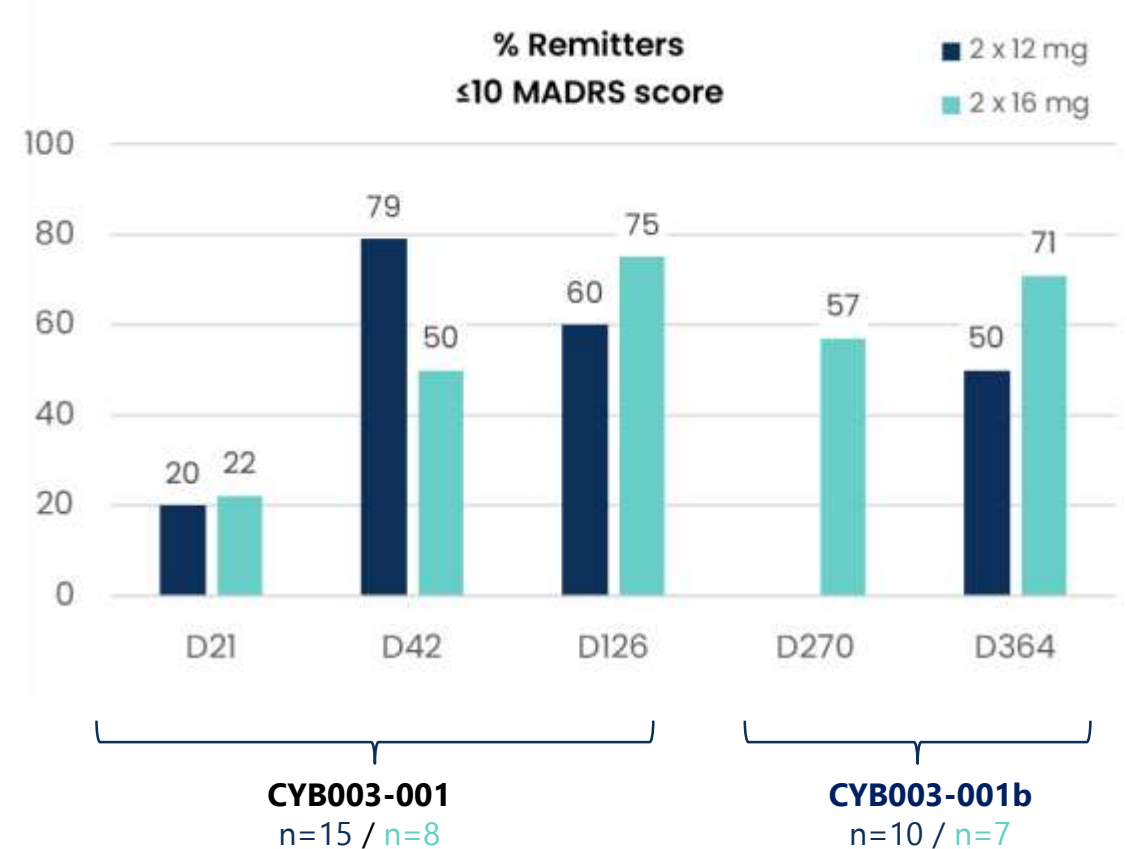
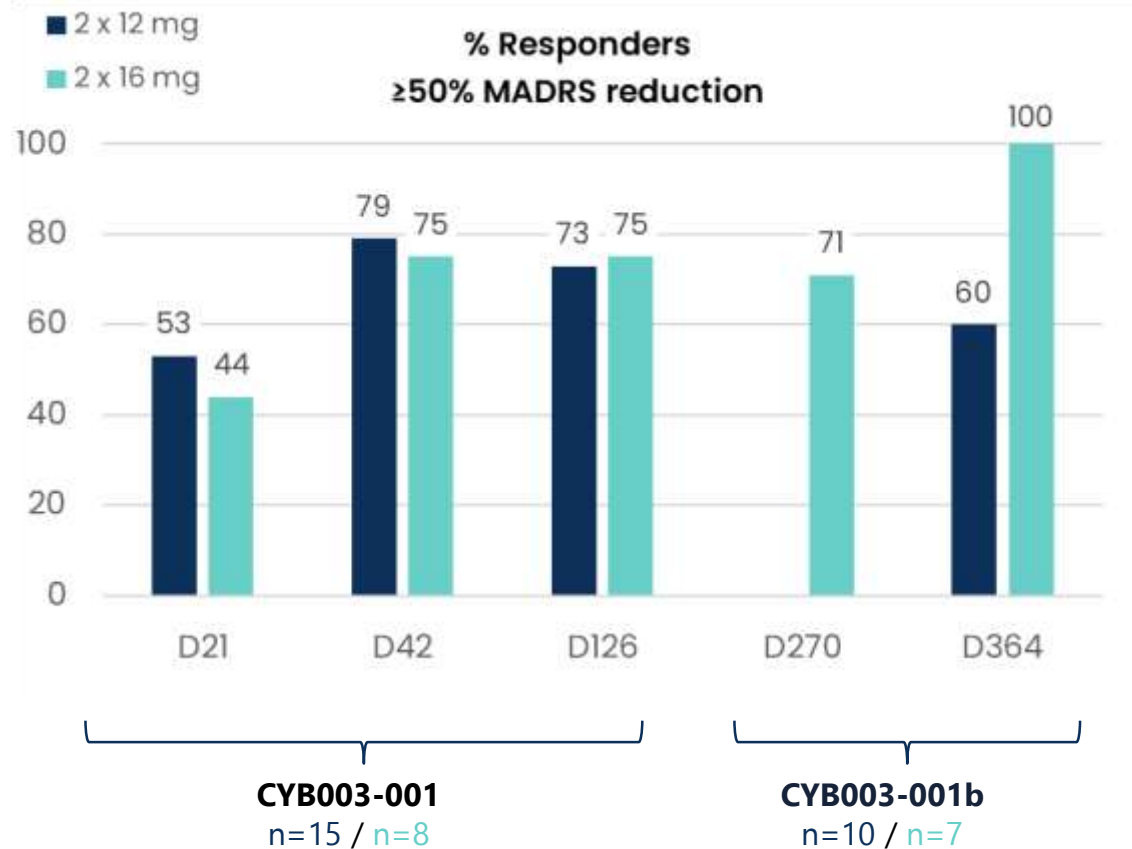
\*Primary endpoint: difference in change from baseline in MADRS total score between CYB003 and placebo at 3 weeks

# Robust, Sustained Improvements in Depression Symptoms at 12 Months

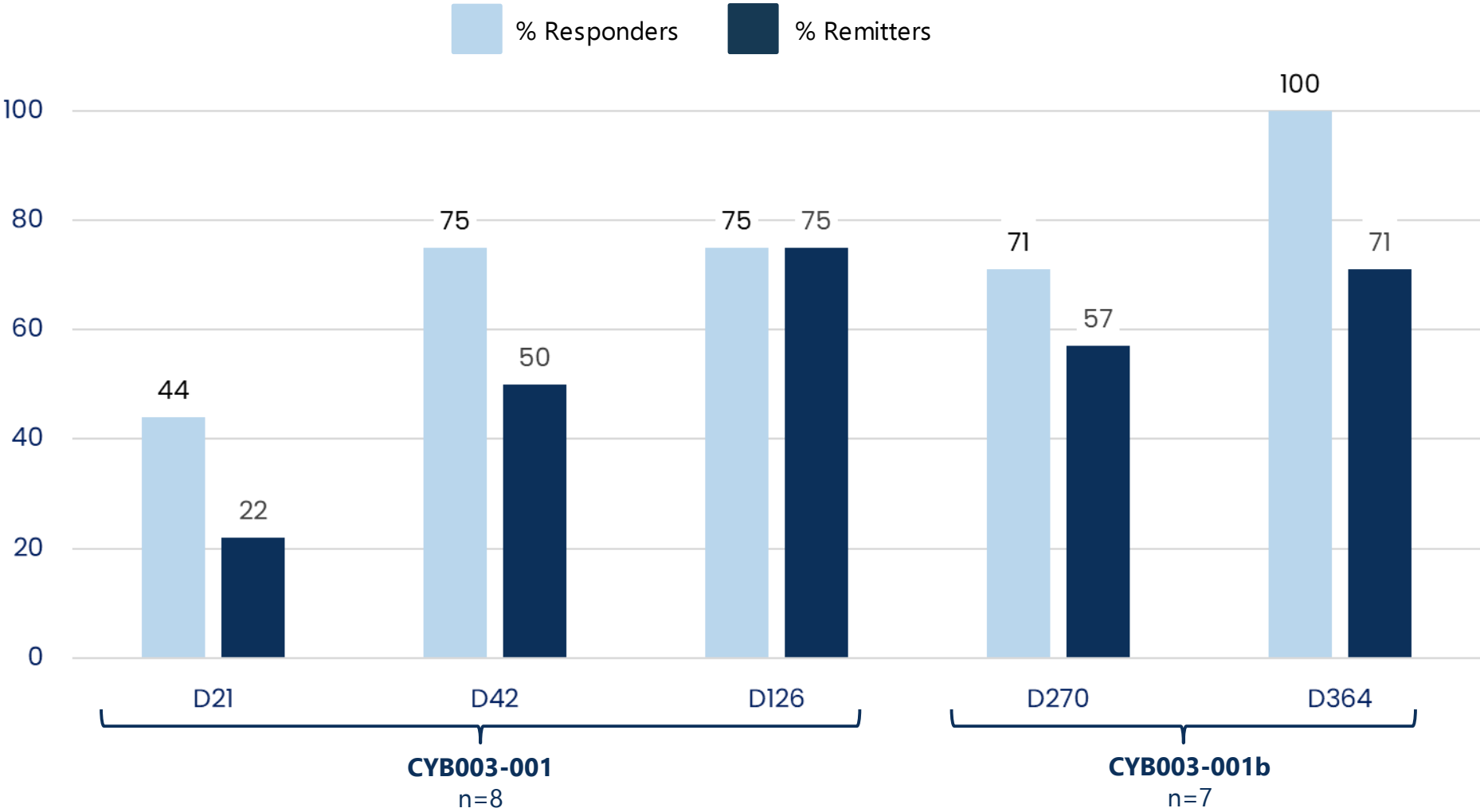




# Response and Remission at 12 Months: 12 mg & 16 mg



# Response and Remission at 12 Months: 16 mg – 2 Doses



# Favorable Safety Profile of CYB003

- No AEs were reported at the 12-month follow up
- No reports of suicidal ideation or behavior or any long-term adverse sequelae

In the short-term study:

- No SAEs and no participant discontinued the study due to an AE
- Most common AEs were nausea, elevated blood pressure and headache
- Increases in blood pressure were transient and resolved without intervention
- No clinically relevant changes in chemistry, hematology markers or ECG parameters

# 12-Month Data Highlights: Phase 2 CYB003 Results in MDD

## Robust, sustained improvements in depression symptoms



- Mean ~23-point reduction in MADRS scores from baseline at 12 months following 2 doses (16mg)
- Average baseline MADRS score was ~32

## Durable response and remission rates



12 months after 2 doses (16 mg):

- 100% of patients were responders
- 71% of patients were in remission

## Favorable safety and tolerability profile



- All reported AEs mild to moderate; no AEs of suicidality
- No AEs/SAEs reported in the 12 month follow up

# Phase 3 PARADIGM Program Overview

Study design aligned with FDA guidance and two meetings with FDA

Addressing functional unblinding

Phase 3 underway

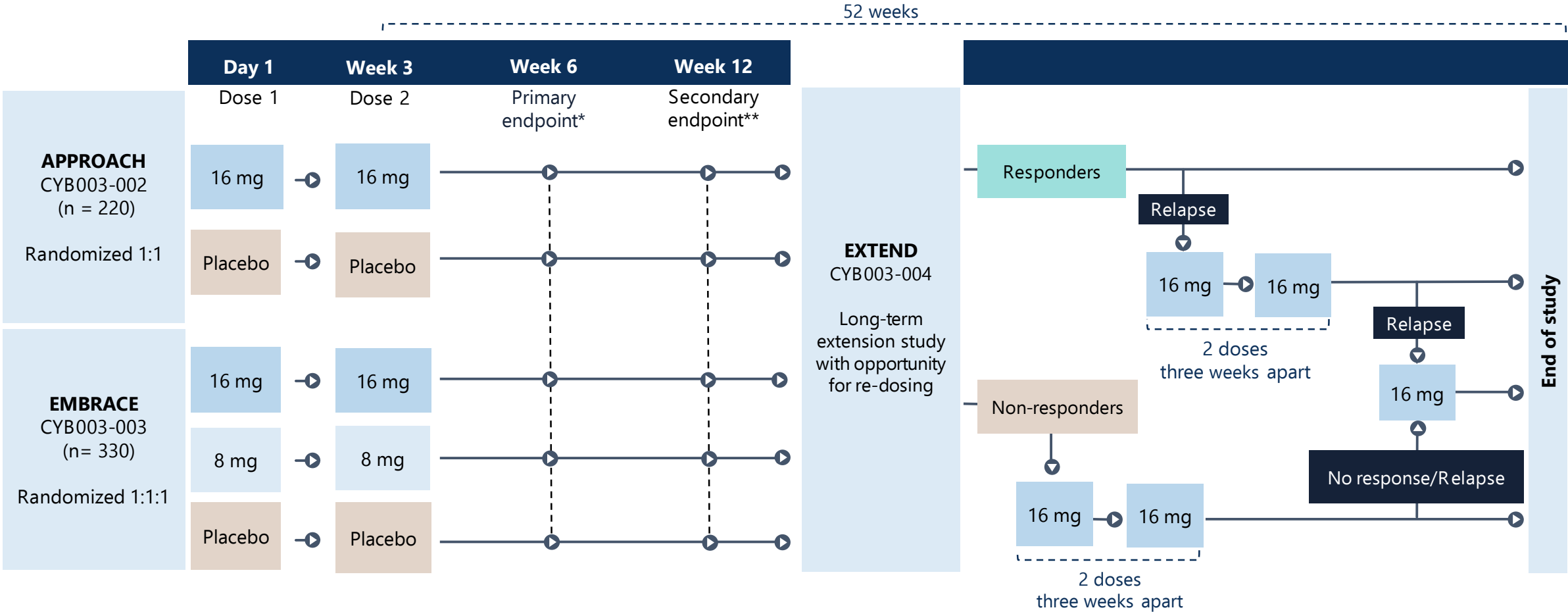
The pivotal program will consist of 2 studies plus an extension<sup>1,2,3</sup>:

- APPROACH: Two-arm study of two doses of CYB003 vs. placebo
- EMBRACE: Three-arm study with a high dose, mid-dose, and placebo arm
- EXTEND: Long-term extension study that allows for open-label dosing or re-dosing for participants who did not respond in the first two studies or relapsed during the extension study
- Use of remote, independent, blinded raters
- Firewall effects reporting during the dosing session
- Long-term efficacy data points up to one year to outlast expectancy bias
- Multinational Phase 3 program will include more than 45 sites across 12 countries in the U.S. and Europe<sup>1,2,3</sup>
- Study sites selected with clinical expertise and training in depression studies
- Clinical supplies manufactured and ready

Notes:

- 1) Forward-looking statements are subject to risks and assumptions. See "Cautionary Statement" on page 2 of this presentation.
- 2) Subject to receipt of all necessary regulatory approvals from all applicable governmental authorities, including, as applicable, the academic and scientific organizations with which Cybin is working. There are multiple risk factors regarding the ability to successfully commercially scale a chemically synthesized process to obtain psilocin and other analogues. There is no assurance that timelines will be met. Anticipated timelines regarding drug development are based on reasonable assumptions informed by current knowledge and information available to Cybin. Such statements are informed by, among other things, regulatory guidelines for developing a drug with timeline safety studies, proof of concept studies, and pivotal studies for new drug application submission and approval, and assumes the success of implementation and results of such studies on timelines indicated as possible by such guidelines, other industry examples, and Cybin's development efforts to date.
- 3) Cybin is prioritizing the progression of its CYB003 program. The advancement of Cybin's CYB004, CYB005 and technology programs are all contingent on Cybin's ability to continue raising capital under its current and future financing arrangements. No assurances can be given that Cybin will be able to raise the additional capital that it may require for its anticipated future development.

# PARADIGM: CYB003 Phase 3 Pivotal Program in MDD\*\*\*



\*Primary endpoint: MADRS change from baseline at 6 weeks

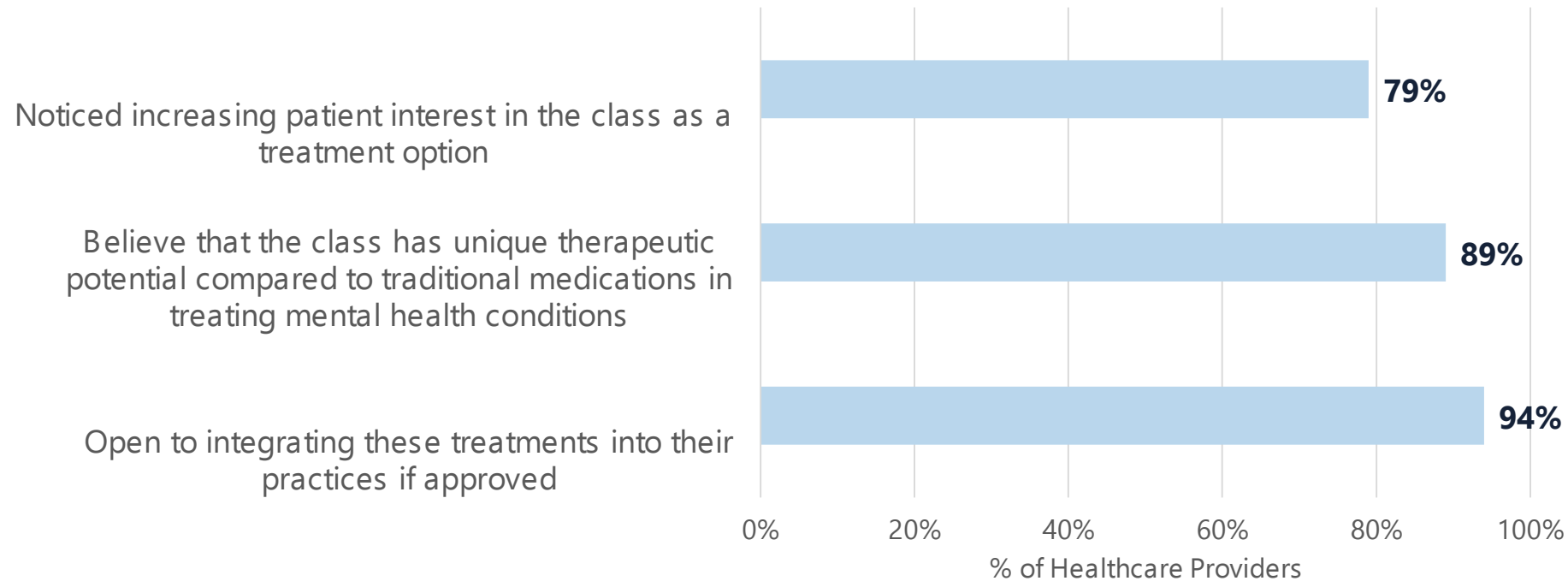
\*\*Key secondary endpoint: MADRS change from baseline at 12 weeks

\*\*\*Note: Forward-looking statements are subject to risks and assumptions. There is no assurance that timelines will be met. Anticipated timelines regarding drug development are based on reasonable assumptions informed by current knowledge and information available to the Company. Such statements are informed by, among other things, regulatory guidelines for developing a drug with safety studies, proof of concept studies, and pivotal studies for new drug application submission and approval, and assume the success of implementation and results of such studies on timelines indicated as possible by such guidelines, other industry examples, and the Company's development efforts to date.




# Increased Awareness Among Healthcare Providers

In a survey of 430+ mental healthcare providers around the world:

## Healthcare Provider Survey Results



# Early Commercial Stakeholder Engagement Underway

Healthcare Providers & Patients 	Clinics 	Payers 
<ul style="list-style-type: none"><li>• Prepare and educate</li><li>• Understand and overcome potential barriers</li></ul>	<ul style="list-style-type: none"><li>• Dosing room setup</li><li>• Site and monitor education and training</li><li>• REMS (Risk Evaluation and Mitigation Strategy) preparedness</li><li>• Explore additional channels beyond existing centers</li></ul>	<ul style="list-style-type: none"><li>• Data collection</li><li>• Pharmacoeconomics</li><li>• Reimbursement route optimization</li></ul>





# CYB004

Deuterated Dimethyltryptamine (dDMT) Program  
Generalized Anxiety Disorder

# CYB004 Program Overview

## Short-duration treatment with convenient dosing

- Short-duration treatment
- Intramuscular dosing is more convenient and patient-friendly vs. IV

## Demonstrated proof-of-concept in depression and anxiety

- Strong datasets across 5 studies supporting characterization and dosing optimization for dDMT
- Robust efficacy in depression with improvements in anxiety scores
- Favorable safety profile

## Robust IP Protection for DMT/dDMT

- >50 patents in support of CYB004 program including composition of matter protection

# Target Product Profile for dDMT

## Optimized with Data from 5 Clinical Studies

### Completed Studies

- 1 Phase 1/2a DMT study in moderate to severe MDD (no SSRIs)
- 2 Phase 1 IV/IM DMT study
- 3 Phase 1 SSRI DDI study
- 4 Phase 1 Study of IV CYB004 (dDMT) and IV DMT
- 5 Phase 1 IM/IV dDMT study



### Key Findings

#### **Rapid and durable antidepressant and anxiolytic effect observed in DMT**

- ✓ 47% of MDD patients in remission at 3 months
- ✓ Among the patients that achieved remission at 3 months, 64% had sustained remission at 6 months
- ✓ 40% of MDD patients in remission at 6 months
- ✓ Rapid improvement in anxiety and wellbeing scores
- ✓ IV DMT safe and well-tolerated

#### **Characterized safe and well-tolerated IM route and dose selection for DMT and dDMT**

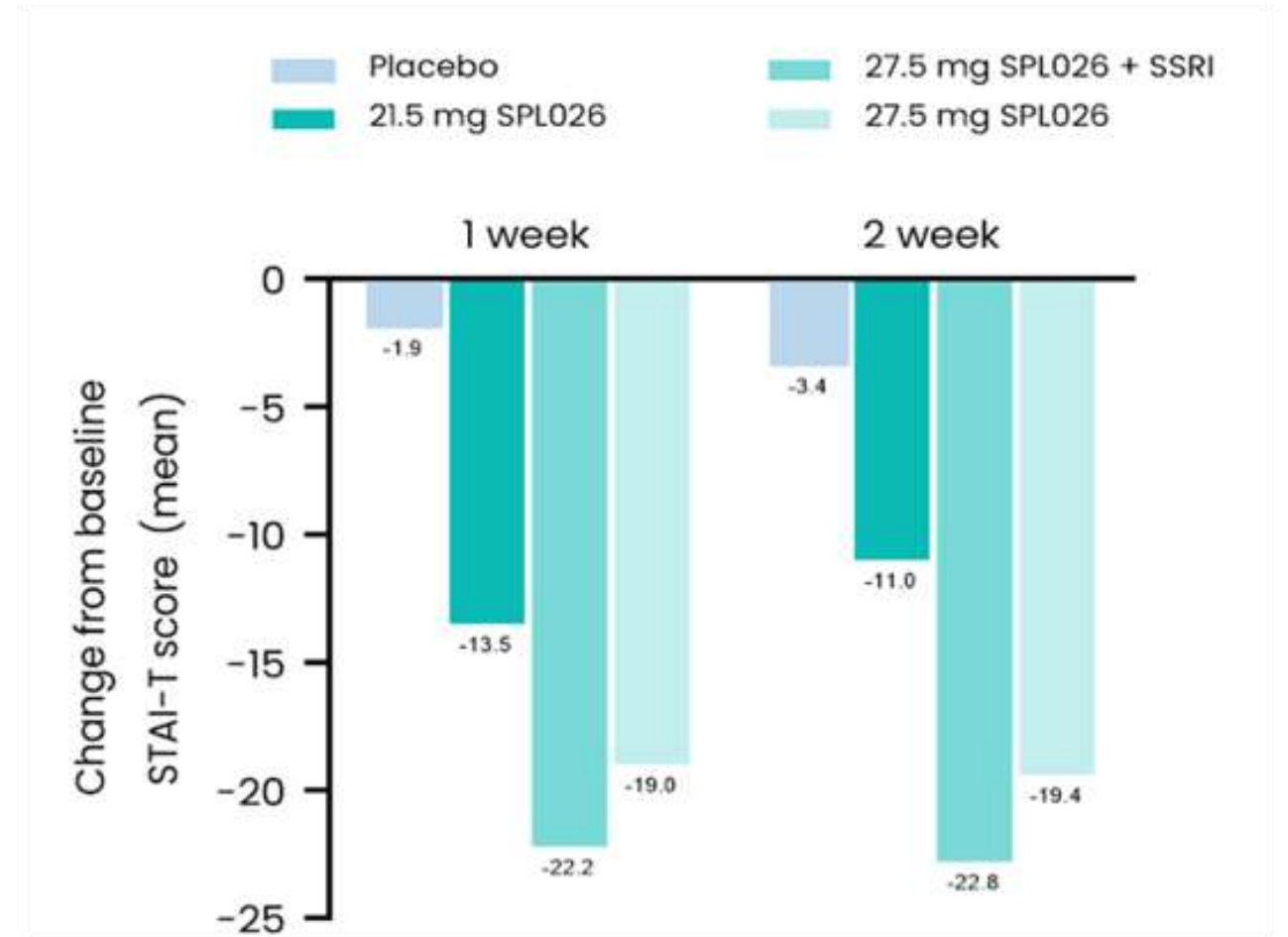
#### **DMT safe and well-tolerated when co-administered with SSRIs**

#### **Potential enhanced effect when given as adjunctive to SSRIs:**

- ✓ 92% remission rate in SSRI cohort vs. 20% remission (non-SSRI cohort)

# DMT Demonstrates Proof-of-Concept in Reducing Anxiety Symptoms<sup>1,2</sup>

- ✓ Efficacy assessed as change from baseline in STAI-T scores
- ✓ Data from the MDD monotherapy (21.5 mg) and SSRI add on studies (27.5 mg)
- ✓ Observed to provide proximal de-risking of development in anxiety

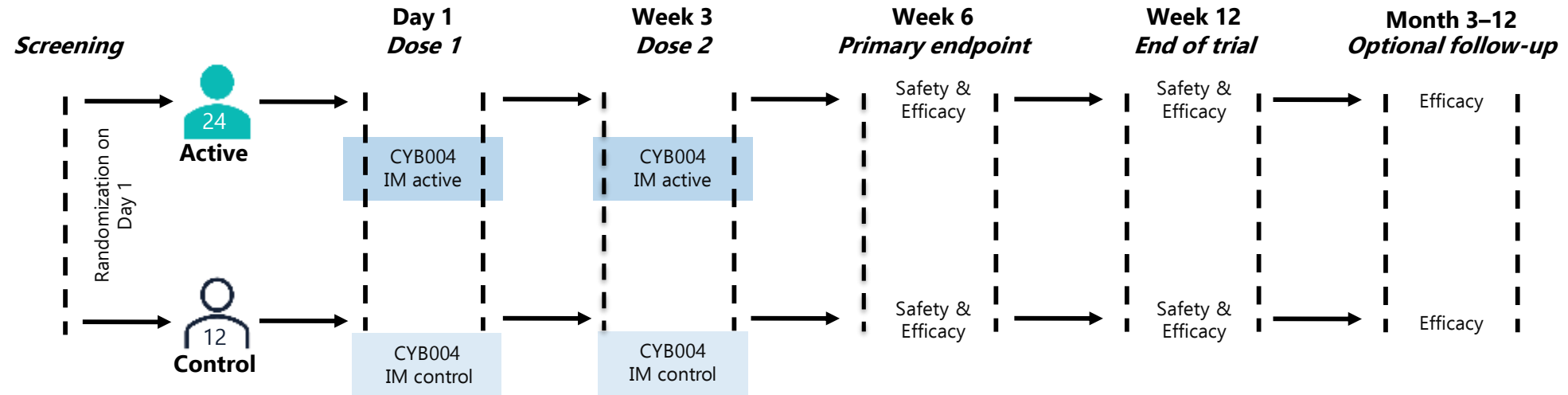


Notes:

1) STAI: State-Trait Anxiety Inventory

2) Doses: 21.5 mg and 27.5 mg doses administered at different rates. 21.5 mg in the Phase 2a MDD study, 27.5 mg in the SSRI DDI study. Placebo data reported is from the Phase 2a study in MDD

# CYB004 in GAD: Phase 2 Proof-of-Concept Study<sup>1</sup>

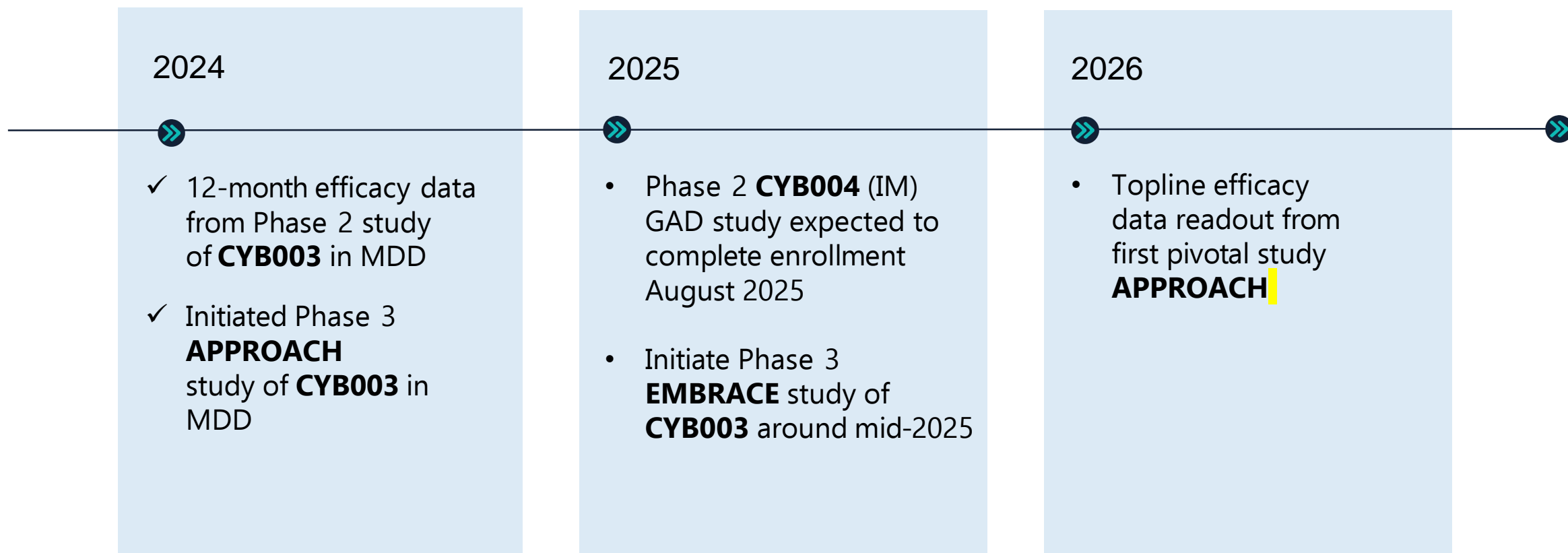


- Moderate to severe GAD
- Concomitant antidepressant/anxiolytic treatment and co-morbid depression allowed

- Primary endpoint: HAM-A
- Other endpoints: HAM-D, safety, EQ-5D-5L

Phase 2 study dosing underway

# Value-Driving Milestones Across Development Pipeline<sup>1,2</sup>



# Thank You

NYSE American: CYBN  
Cboe CA: CYBN

Contact: [ir@cybin.com](mailto:ir@cybin.com)

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- \*Up to 37% suffer from nausea, diarrhea, constipation, vomiting, dry mouth, and rarely gastrointestinal bleeding (based on a review of package inserts)

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