

Transforming Mental Health Care

Investor Presentation
November 2024



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Compass Pathways

Dedicated to accelerating patient access to evidence-based innovation in mental health.

- Lead product candidate: COMP360 psilocybin treatment in treatment resistant depression (TRD)
- Phase 2 TRD program published in *The New England Journal of Medicine*
Phase 3 TRD program recruiting
 - Pivotal trial 1 (COMP005): top-line data expected Q2 2025
 - Pivotal trial 2 (COMP006): top-line expected H2 2026
- Phase 2 PTSD positive top-line data reported in Q2 2024



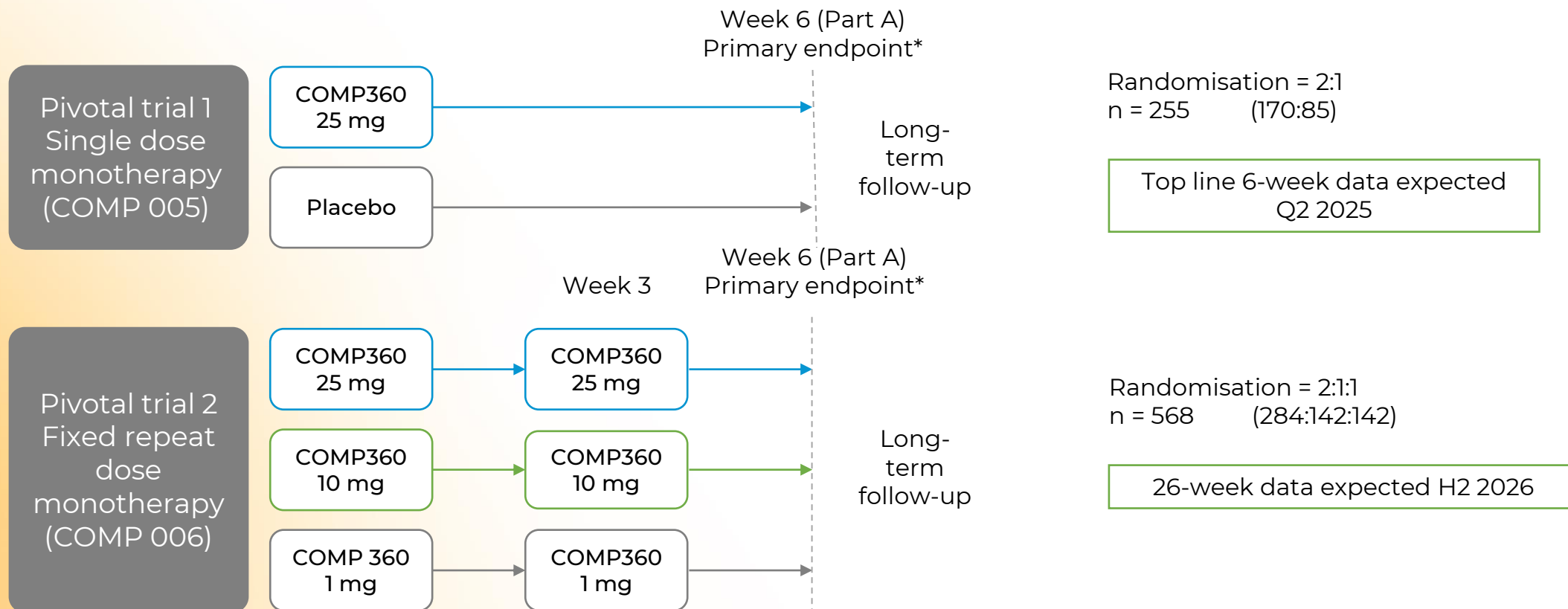
TRD treatment pathway: significant unmet need for 100 million patients

Treatment pathway stage	New onset depression Major depressive disorder (MDD)	Persistent depression Major depressive disorder (MDD)	Treatment-resistant depression (TRD)
Line of therapy	First line	Second line	Third line + 
Estimated number of patients (worldwide)	320 million	200 million	100 million (~1 in 3 of total) US health care cost approx. \$17- 25k per patient/year
Available treatments	<ul style="list-style-type: none"> - Antidepressants - Psychological interventions, e.g., CBT* 	<ul style="list-style-type: none"> - Antidepressants - Antidepressant combinations - Psychological interventions 	<ul style="list-style-type: none"> - Antidepressants - Augmentation therapy (antidepressants, mood stabilizers, anticonvulsants, atypical antipsychotics, esketamine) - Ketamine - Somatic therapy (rTMS, tDCS, ECT, DBS)* - High-intensity psychological interventions
% relapse	60-70%	50-75%	80-90%

*NOTE: CBT = cognitive behavioural therapy; rTMS = repetitive transcranial magnetic stimulation; tDCS=transcranial direct current stimulation; ECT=electroconvulsive therapy; DBS=deep brain stimulation
SOURCE Table adapted from Rush, A. J., Trivedi, M. H., Wisniewski, S. R., Nierenberg, A. A., Stewart, J. W., Warden, D., ... & Fava, M. (2006). Acute and longer-term outcomes in depressed outpatients requiring one or several treatment steps: a STAR* D report. American Journal of Psychiatry, 163(11), 1905-1917; Zhdanova M, Pilon D, Ghelerter I, et al. The prevalence and national burden of treatment-resistant depression and major depressive disorder in the United States. J Clin Psychiatry. 2021;82(2):20m13699.



Phase 3 program: Overview of ongoing pivotal trial designs

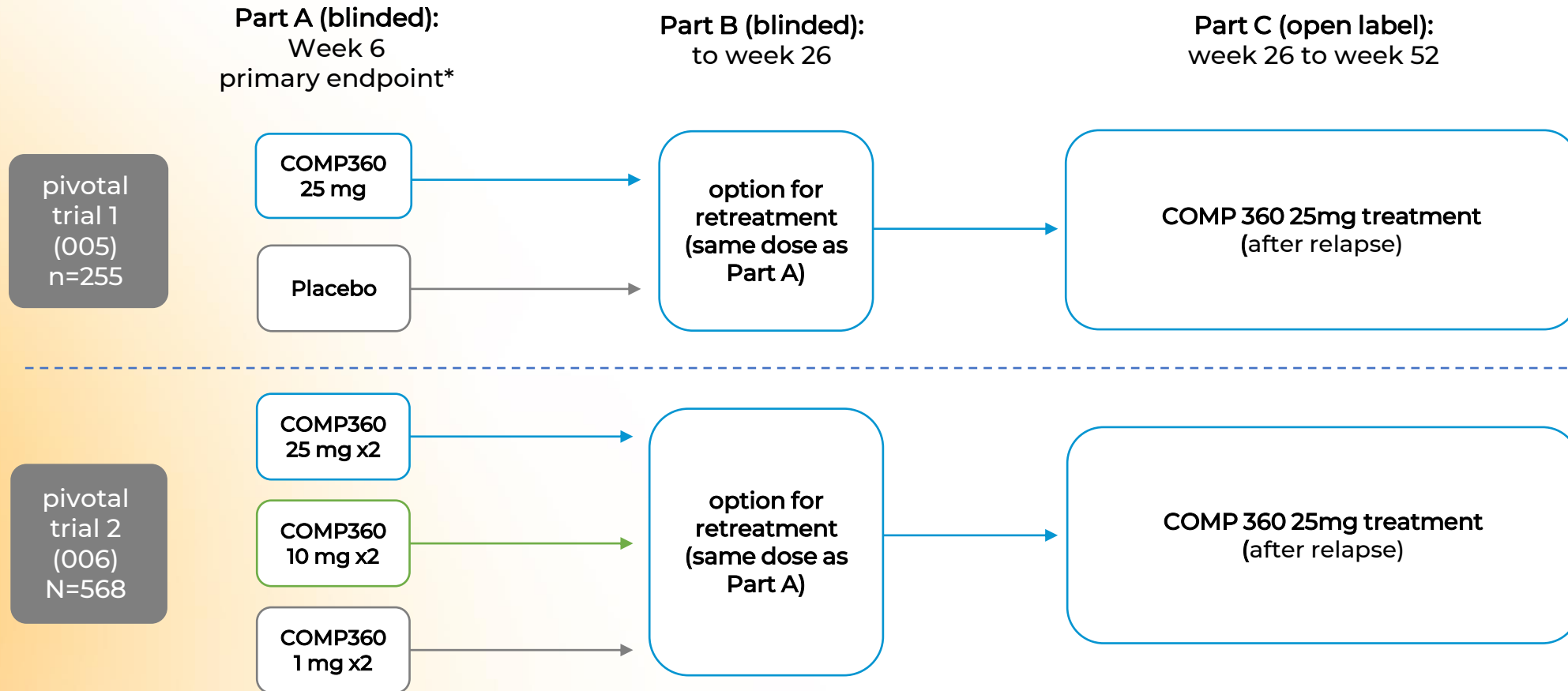


The phase 3 program will be conducted across approx. 150 sites in 12 countries. The participant population (TRD definition and core inclusion/exclusion criteria) remains unchanged compared to Phase 2b

*Primary endpoint - change from baseline in MADRS total score at Week 6



Phase 3 program long-term follow up component



*Primary endpoint - change from baseline in MADRS total score at Week 6



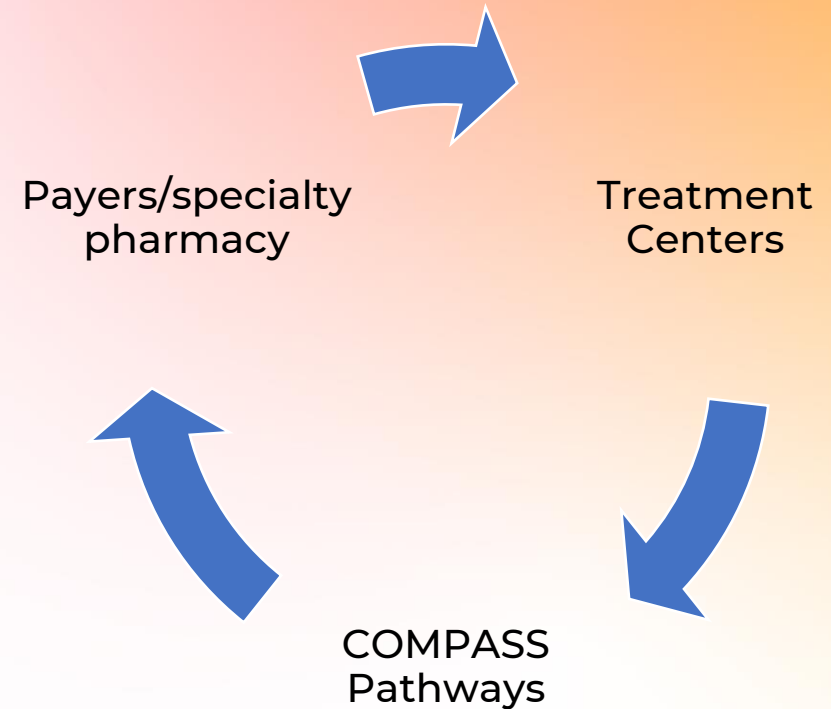
Preparing for scale at launch, if approved

Compass Pathways intends to deliver COMP360 (medicine) to **Treatment Centers** through specialty pharmacy channels, reimbursed by **Payers**

Our strategy for reimbursement is for **Treatment Centers** to be reimbursed by **Payers** with new reimbursement codes specific to psychedelic therapies*

Strategic research collaborations established to inform the development of scalable and practical delivery models for COMP360 psilocybin treatment, if approved for treatment-resistant depression

Regulatory approval and payer coverage/reimbursement is the path to broad and equitable patient access



*New CPT III codes accepted, and language released by AMA for Psychedelic Drug Monitoring Services, expected to be published in the CPT Manual and become effective on January 1, 2024



The infrastructure to deliver COMP360 psilocybin treatment already exists and is growing

Specialty TRD centers, health systems, and integrated delivery networks (IDNs), some of which are clinical trial sites during our phase 3 clinical program, are already experienced in delivering interventional psychiatry treatments like ketamine, Spravato® (esketamine), transcranial magnetic stimulation (TMS) and electroconvulsive therapy (ECT) to tens of thousands of TRD patients

These are delivered relatively frequently, requiring a cumulatively high number of hours of patient and provider time

TMS:
30-36 treatments⁽²⁾⁽⁵⁾

esketamine:
20-28 treatments⁽¹⁾

ketamine:
12-15 treatments⁽⁶⁾

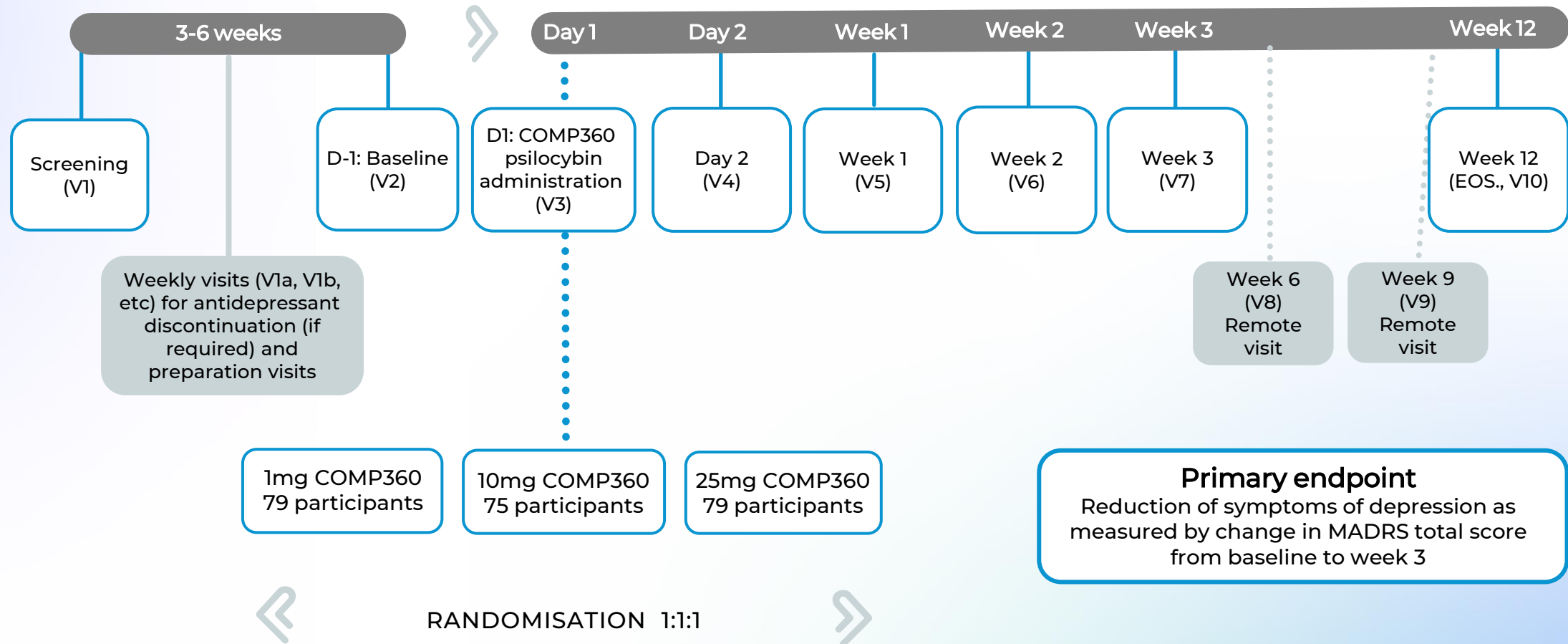
ECT:
6-12+ treatments⁽³⁾⁽⁴⁾

*treatment #s represent a typical course over 6 months

References: [1] ICER, 2019; [2] Ross, 2018; [3] Petrides, 2011; [4] Thirthalli, 2020; [5] Voigt, 2017; [6] Wilkinson, 2018.



COMP001 phase 2b study design and primary endpoint (n=233)



Note: MADRS = Montgomery-Åsberg Depression Rating Scale; EOS = end of study; TRD = treatment-resistant depression; D = day; V = visit



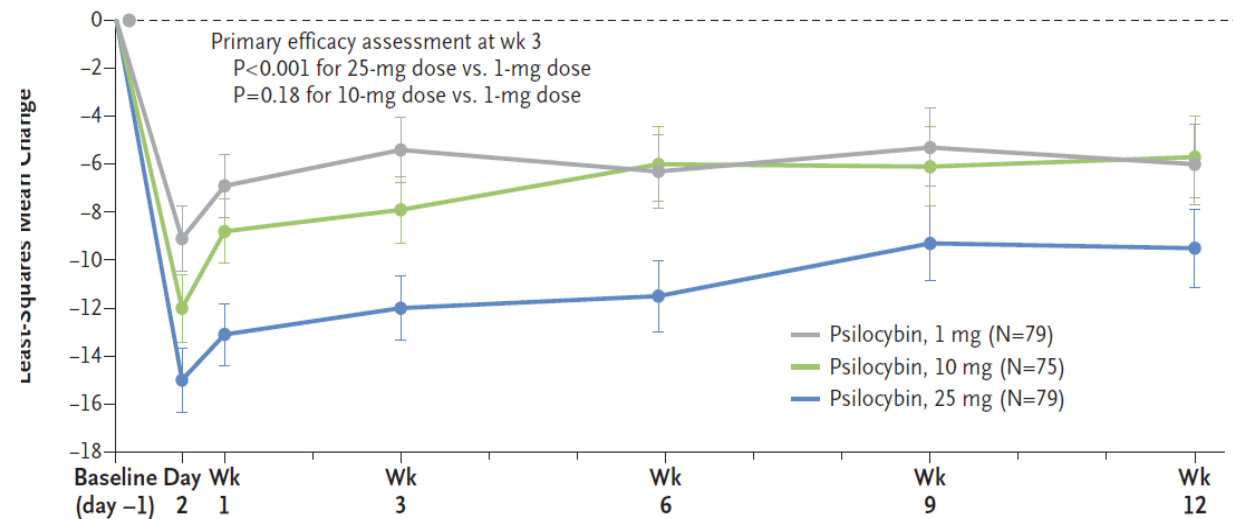
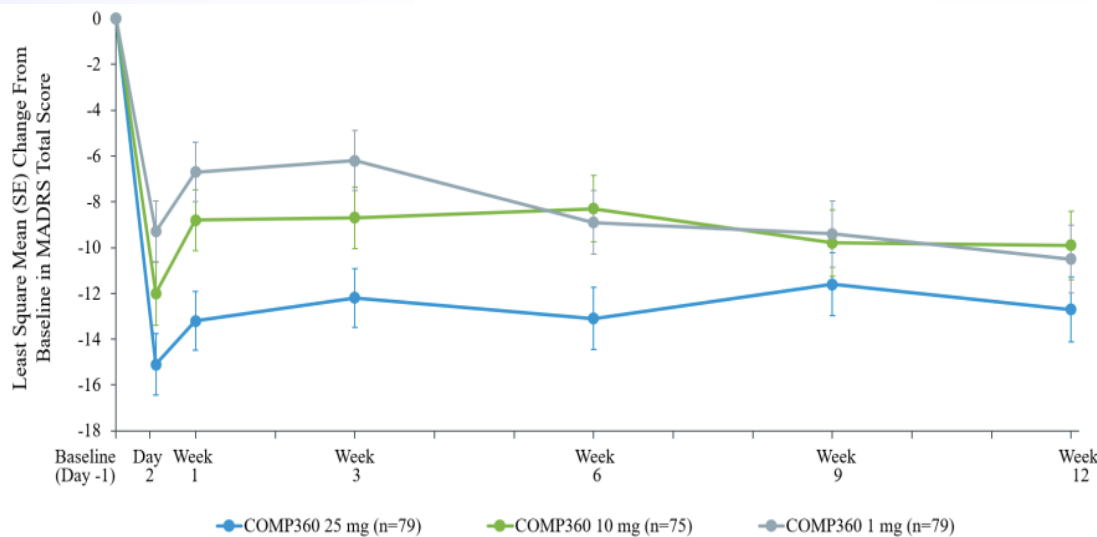
Phase 2b trial: Results demonstrate the potential for a rapid, sustained response in TRD

Published in The NEW ENGLAND JOURNAL of MEDICINE

Randomized, controlled, double-blind trial, 3 arms, single dose (either 1mg, 10mg or 25 mg) of COMP360 psilocybin alongside psychological support. Results were measured as a change on the MADRS* depression scale from baseline (a day prior to administration) over a 12-week period. The primary endpoint of this study was the change from baseline in MADRS total score at week 3.

Without imputation for use of anti-depressants during trial period (consistent with phase 3 design)

With imputation for use of anti-depressants during trial period (applied in phase 2b)



Efficacy: We saw a statistically significant and clinically meaningful reduction in depression symptoms.

Rapid onset of action: The effect occurred the day after the administration.

Durability: We saw a sustained response at week 12 – a positive indication for high potential as a monotherapy.



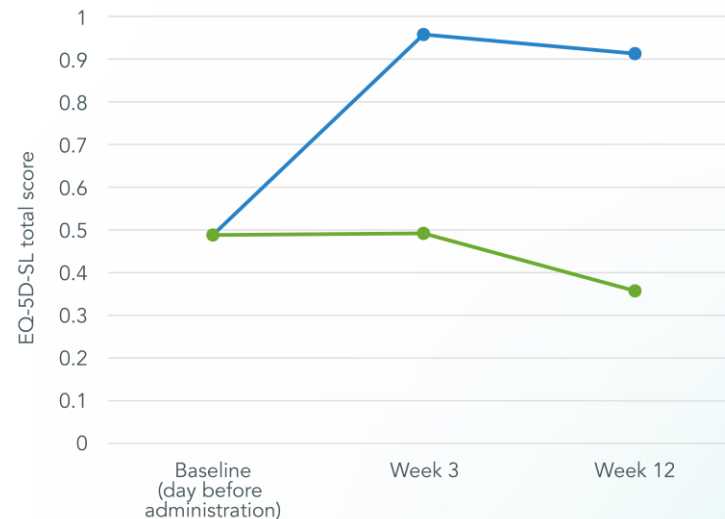
Phase 2b trial: Those participants who showed a sustained response also showed signs of improvement beyond the reduction of depression symptoms

Sustained responders are participants who responded ($\geq 50\%$ change in MADRS total score from baseline) at weeks 3 and 12, and at least one visit out of week 6 and 9, and who did not start new treatments for depression.

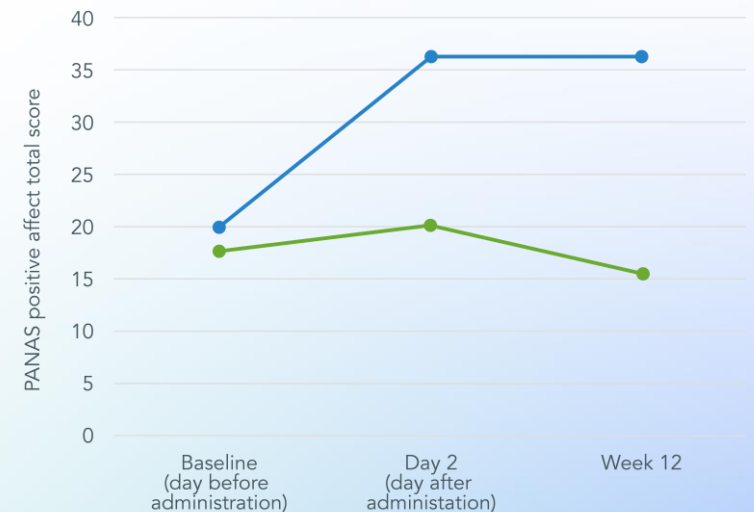
Sustained non-responders are participants who did not respond ($< 25\%$ change in MADRS total score from baseline) at weeks 3 and 12, and at least one visit out of week 6 or 9.

- Sustained responders (n=19)
- Sustained non-responders (n=21)

Quality of life: Sustained responders were found to have a clinically meaningful increase in quality of life from baseline at week 3 and week 12 with scores in the normal range after treatment



Positive affect: Sustained responders were found to have a clinically meaningful increase in positive affect from baseline on the day after the psilocybin session and at week 3



NOTE: EQ-5D-3L= EuroQoL 5-Dimensions 3-Levels; PANAS= Positive and Negative Affect Schedule; SD= standard deviation



Phase 2b trial: COMP360 psilocybin treatment was generally well-tolerated

Treatment-emergent adverse events (TEAEs)

>90% of TEAEs were of mild or moderate severity.

5

most frequent TEAEs across the 10mg and 25mg doses were headaches, nausea, fatigue, insomnia and anxiety.

>77%

of TEAEs occurring on the day of administration resolved on the same or next day; most were mild or moderate.

There were no concerns with vital signs, ECG or clinical laboratory data in any of the treatment groups

TEAEs involving hallucinations (which only occurred in the 25mg and 10mg groups) and illusions (all groups) started and resolved on the day of administration.

TESAEs of suicidal ideation, suicidal behaviour and intentional self-injury were uncommon but occurred unevenly across groups in non-responders

- All patients who experienced these events during the trial had said during screening that they had had suicidal thoughts prior to the trial.
- 3 TESAEs of suicidal behavior in non-responders, 30 days post administration in the 25 mg arm emphasizing the need for a vigilant approach to the TRD condition.



Most frequent TEAEs ordered by the 25mg arm (at least 5% in any treatment group)

MedDRA TEAE preferred term	COMP360 25mg	COMP360 10mg	COMP360 1mg	Overall
	N=79	N=75	N=79	N=233
	n (%)			
Headache	27 (34.2)	16 (21.3)	20 (25.3)	63 (27.0)
Nausea	18 (22.8)	7 (9.3)	4 (5.1)	29 (12.4)
Fatigue	12 (15.2)	5 (6.7)	7 (8.9)	24 (10.3)
Insomnia	8 (10.1)	11 (14.7)	14 (17.7)	33 (14.2)
Anxiety	7 (8.9)	13 (17.3)	3 (3.8)	23 (9.9)
Mood altered	7 (8.9)	3 (4.0)	1 (1.3)	11 (4.7)
Back pain	6 (7.6)	0	3 (3.8)	9 (3.9)
Dizziness	6 (7.6)	1 (1.3)	1 (1.3)	8 (3.4)
Suicidal ideation	5 (6.3)	5 (6.7)	4 (5.1)	14 (6.0)
Myalgia	5 (6.3)	2 (2.7)	1 (1.3)	8 (3.4)
Euphoric mood	4 (5.1)	5 (6.7)	4 (5.1)	13 (5.6)
Depression	4 (5.1)	6 (8.0)	5 (6.3)	15 (6.4)
Abdominal pain upper	4 (5.1)	2 (2.7)	1 (1.3)	7 (3.0)
Irritability	4 (5.1)	2 (2.7)	1 (1.3)	7 (3.0)
Panic reaction	4 (5.1)	1 (1.3)	1 (1.3)	6 (2.6)
Depressed mood	3 (3.8)	5 (6.7)	4 (5.1)	12 (5.2)
Paraesthesia	3 (3.8)	4 (5.3)	1 (1.3)	8 (3.4)
Thinking abnormal	0	4 (5.3)	0	4 (1.7)

TEAE incidence is higher in the 25mg group overall

Key mood-related TEAEs (euphoric mood, depression, depressed mood, suicidal ideation) do not have a higher incidence in the 25mg arm

Note: MedDRA = Medical Dictionary for Regulatory Activities; TEAE = treatment emergent adverse event; N = number of participants in the population; n = number observed

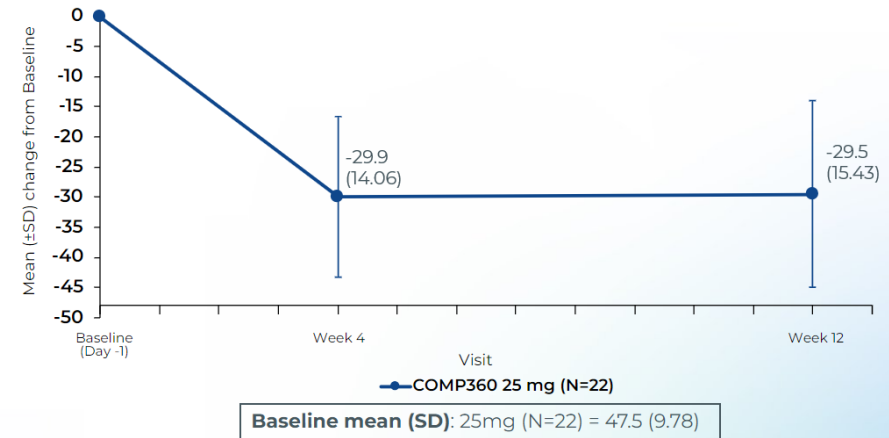


Post-traumatic stress disorder (PTSD) positive phase 2 study

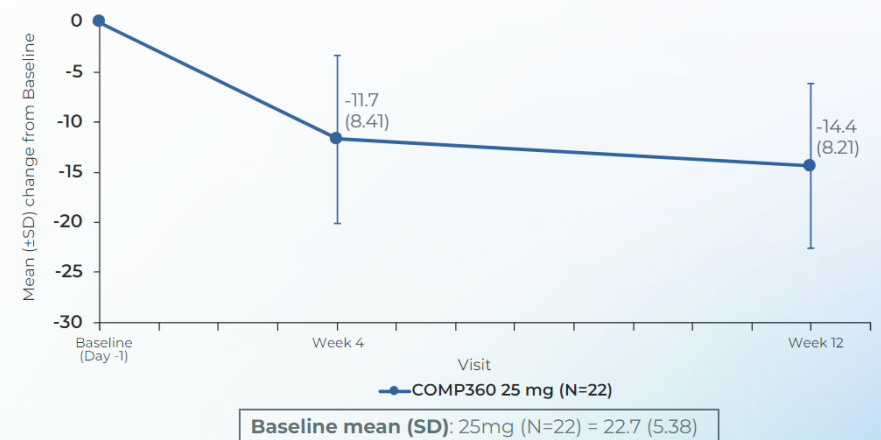
- N=22, multi-center open-label, single administration of 25mg COMP360 with psychological support
- Early onset and sustained change from baseline in CAPS-5 observed at week 4 and week 12
- Durability in CAPS-5 reductions from baseline seen at week 4 (29.5 points) and week 12 (29.9)
- Response in CAPS-5: 81.8% at week 4, 77.3% at week 12
- Remission in CAPS-5: 63.6% at week 4, 54.5% at week 12
- No treatment emergent serious adverse events reported
- Mean baseline of 47.5 CAPS-5 total score, which is considered severe

NOTE: CAPS-5 = clinician administered PTSD scale

Summary of change from baseline in CAPS-5 score



Summary of change from baseline in SDS score



Phase 2 PTSD study safety profile (primary endpoint)

Summary of most frequent TEAES ($\geq 10\%$ prevalence)

COMP360 was well tolerated with no treatment emergent serious adverse events reported

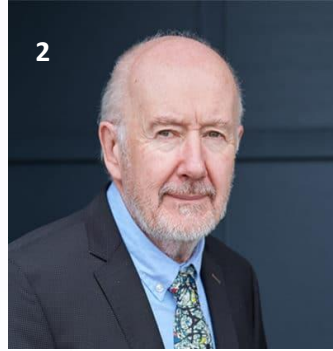
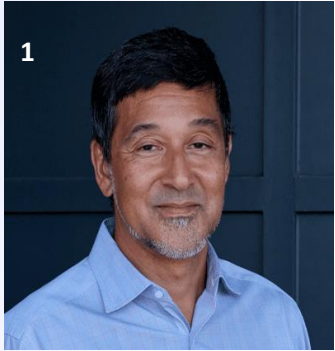
No participants re-started SSRI's or antidepressants after COMP360 administration in study

MedDRA TEAE Preferred Term (at least 5%)	COMP360 25 mg (N = 22)			
	Overall		COMP360 admin day	
	n (%)	E	n (%)	E
Headache (PTs: Headache, Tension headache)	11 (50.0)	15	6 (27.3)	6
Nausea	8 (36.4)	9	6 (27.3)	6
Crying	6 (27.3)	6	6 (27.3)	6
Fatigue	6 (27.3)	6	4 (18.2)	4
Hallucination (PTs: visual, auditory, synaesthetic)	5 (22.7)	7	5 (22.7)	7
Muscle tightness	3 (13.6)	3	3 (13.6)	3
Paraesthesia	3 (13.6)	3	2 (9.1)	2
Visual impairment	3 (13.6)	3	3 (13.6)	3

NOTE: E = events, MedDRA = Medical Dictionary of Regulatory Authorities, n = number of participants with TEAE, PT = preferred term, TEAE = treatment emergent adverse event



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COMPASS runway expected into 2026

Cash and cash equivalents at September 30, 2024

\$207 million

Financial guidance for cash used in operating activities (As issued on Q3 2024 earnings call on 31 October 2024)

Fourth quarter 2024:

\$37 - \$43 million

Full-year 2024:

\$114 - \$120 million



We're a biotechnology company...

...dedicated to accelerating patient access to evidence-based innovation in mental health.

Stephen Schultz
SVP, Investor Relations
stephen.schultz@compasspathways.com
+1 401-290-7324

