

Everyone deserves the option of better mental health

TSXV: DMT

OTCQB: DMTTF

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Introducing Small Pharma^{a,b,c}

- Depression represents a huge unmet need with 280 million people suffering globally (2019)¹. Only one-third of patients respond to existing first line treatments², and many struggle with side effects³.
- Small Pharma is developing novel and protectable psychedelic-based therapies that have the potential to offer meaningful mental health and broader wellbeing improvements for patients.
- We target **short in-clinic treatment sessions** that last <2.5 hours, enhancing clinical convenience relative to alternative psychedelic treatments in development, such as psilocybin and LSD, that typically last a full day in clinic.^{4,5}
- Clear proof-of-concept for DMT-based therapy in treating major depression as demonstrated through results from the Company's SPL026 Phase IIa study, with rapid-acting and long-lasting antidepressant effects to at least 6 months.⁶
- Advancing portfolio with positive progress of second clinical candidate, SPL028 a 2nd generation deuterated DMT asset with robust IP protection, anticipated to offer a differentiated DMT treatment profile.⁷

There is a huge unmet need in depression

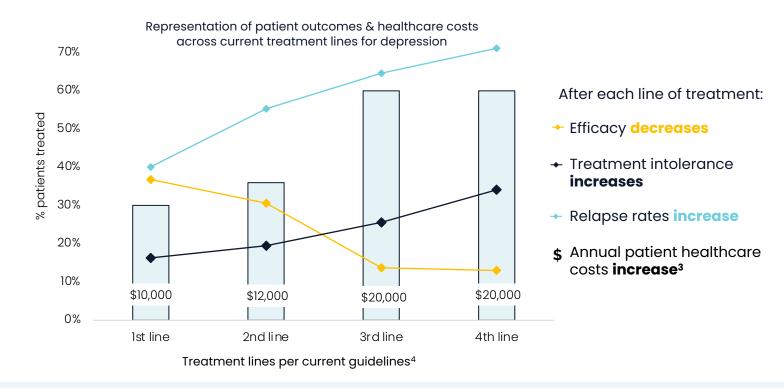
Unmet Opportunity

Depression is a chronic illness that affects people everywhere.

- ~280m people suffer worldwide¹
- Costs global economy \$1 trillion each year in lost productivity¹
- Suicide risk is 20x times higher for an individual with vs. without depression²
- #1 leading cause of disability worldwide1

Unmet Care

Current treatment options leave millions of patients behind. 1st line treatments work for approximately one-third of patients. The treatment success rate decreases at each successive treatment line, as patients try to find one that works for them.



a-c: See Appendix - Footnotes and Sources and "Cautionary Notes - Forward-Looking Information", "Risk Factors" and "Treatment Claims"

Corporate Presentation | July 2023

Our portfolio targets scalable in-clinic DMT-based treatments with differentiated clinical & commercial benefits a.b.c

Anticipated treatment journey

INITIAL CONSULTATION

- Evaluate eligibility
- Preparation and education provided to patient



DOSING & PSYCHEDELIC EXPERIENCE 30 MINS - 1 HR1

In-clinic treatment

PREPARATION

15-30 MIN

- Episodic, as required dosing regimen
- Supervised by licensed practitioner

INTEGRATION **TALK THERAPY**

1 HR+

· Ongoing assessment to evaluate re-treatment needs

Follow up psychological

FOLLOW UP

support

- delivered by trained therapists to maximize therapeutic outcomes

Target treatment value proposition

RAPID AND DURABLE EFFICACY

WELL-TOLERATED WITH MINIMAL

TREATMENT

Pre-and post dosing psychological support



A neuropsychiatry pipeline of short-duration psychedelics with therapy a,b,c

We leverage data from early-stage trials to inform ongoing portfolio strategy

	PIPELINE CANDIDATE	PRECLINICAL	PHASE 1	PHASE 2A	PHASE 2B	STUDY RATIONALE
First- generation molecule	SPL026 DMT IM or IV	Phase 1/2a in MDD	(IV) - completed			 ✓ Demonstrated proof-of-concept for DMT-based therapies in treating MDD
		Phase 1 IM / IV - c c	ompleted			 ✓ Explore potential alternative administration route (IM vs IV) ✓ Inform pharmacokinetic modelling for SPL028 program
		Phase 1b SSRI Drug Interaction study (((IV) - active			 Assess potential interaction between DMT-based treatment and first-line antidepressants
Second- generation molecule	SPL028 Injectable deuterated DMT	Phase 1 IM & IV - a	ctive			 ✓ Confirm SPL028 meets target drug profile in Phase I program ✓ Option to evaluate efficacy signals in depression patient population via expansion of Ph I program
Second- generation molecule	SPL029 Oral tryptamine series	Preclinical				✓ Candidate selection ongoing



6

Clear proof-of-concept achieved for DMT-based therapy a,b,c

Phase IIa trial demonstrated a rapid and durable efficacy profile for IV SPL026 with supportive therapy

PHASE IIA STUDY MET PRIMARY ENDPOINT

Primary endpoint met with a statistically significant -7.4 point difference between SPL026 (21.5mg IV) and placebo at two-weeks post-dose, as measured by MADRS change from baseline (p=0.02)

RAPID & DURABLE ANTIDEPRESSANT EFFECT

- Rapid onset antidepressant effects demonstrated at one-week post-dose with a statistically significant difference in MADRS of -10.8 versus placebo (p=0.002)
- Durable antidepressant effect with a 57% remission rate at 12-weeks following a single SPL026 dose with supportive therapy¹
- No differences identified in antidepressant effect between a one and two dose regimen of SPL026

FAVORABLE SAFETY PROFILE

- SPL026 demonstrated a favorable safety profile and was well-tolerated
- No drug-related Serious Adverse Events (SAEs)
- 47 Adverse Events (AEs) deemed possibly related to treatment, all reported to be mild or moderate, and majority resolved during dosing visit

A multi-layered IP strategy offering protection for our portfolio of assets a,b,c

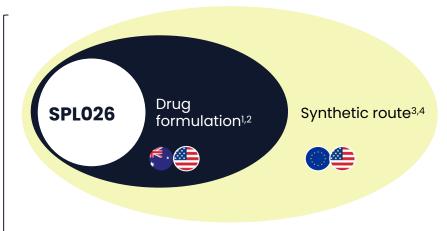
Complete portfolio¹⁵

23 patents granted

patent applications pending

core areas of protection





Medical Related Synthetic Drug **SPL028** COM^{9,10,11,12} use/ COM8 formulation^{1,2,13} route^{3,4} related^{5,6,7,14}

Diagrams represent granted patents surrounding SPL026 & SPL028. Certain granted patents offer protection for both candidates and are illustrated in both diagrams.

5. GB 2 586 940

6. EP 3 902 541

^{2.} US 11 406 619 EP 3 873 883

^{7.} US 11 471 417 9. GB 2 585 978 10. GB 2 592 822

^{11.} EP 3 826 632 12. CA 3 104 072

^{13.} GB 2 595 776 14. JP 7 288 154

Multiple meaningful R&D catalysts expected in next 12 months a,b,c

Cash raised

~C\$63m 2021

Cash position

~C\$18.5m Feb 23¹

Common shares outstanding

321.6m Jun 23²

Completed milestones

- SPL026 Phase 1 HV study
- SPL026 Phase 2a MDD study
- SPL026 Phase 1 IM/IV study
- SPL028 candidate selection
- ☑ Initiation of Phase 1 SPL028 study

H2 2023³

- SPL026 SSRI DDI Ph 1b data
- SPL026 Ph 1 IM/IV data
- SPL028 Ph 1 IM/IV HV data



a-c: See Appendix - Footnotes and Sources and "Cautionary Notes - Forward-Looking Information", "Risk Factors" and "Treatment Claims"

Refers to latest annual MD&A

R&D programs

SPL026 a,b,c DMT fumarate



Rapid & durable efficacy

Positive proof-of-concept data with antidepressant effects demonstrated in patients with MDD from week one to at least six months¹

Safe and well tolerated

Good safety profile with no treatment-related Serious Adverse Events (SAEs) and good tolerability, with no patients reporting that they regretted the experience¹

Short in-clinic treatment

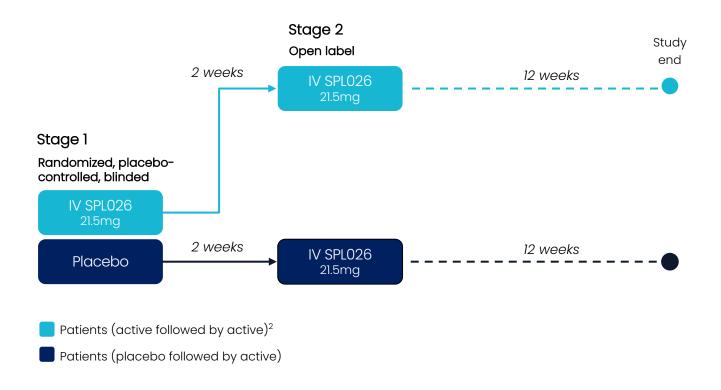
<30 min psychedelic experience demonstrated with IV SPL026¹ and ~45 min demonstrated with IM SPL026², resulting in <2.5 hour treatment session inclusive of supportive therapy</p>

Episodic "as-required" dosing regimen

Based on durability efficacy profile demonstrated in IV SPL026 Phase IIa study, only a few doses may be required in a year dependent on patient need¹

Positive Phase IIa results^{a,b,c}

Robust efficacy profile demonstrated for a DMT-based treatment with supportive therapy in MDD¹



PHASE I

- Psychedelic naïve healthy volunteers (N=32)
- Completed Q3'21
- 21.5mg dose selected as active dose in Phase IIa

PHASE IIA

- MDD patients (moderate/severe) (N=34)
- Not on antidepressant medication/willing to discontinue
- Completed Q4'22

Primary endpoint: blinded phase

- Efficacy: MADRS score change in baseline 2 weeks post dose
- Efficacy also assessed at 1 week post dose

Key secondary endpoint: open label phase

MADRS change from baseline at WI, W2, MI, M3 and M6³ after open label dose

Secondary endpoints: blinded and open label phase

- Safety and tolerability measures
- Assess 1 vs. 2 doses
- Intensity & quality of subjective psychedelic experience measures



a-c: See Appendix - Footnotes and Sources and "Cautionary Notes - Forward-Looking Information", "Risk Factors" and "Treatment Claims" Based on intravenous (IV) SPL026 Phase IIa data suggestive of antidepressant effect as measured using MADRS

^{2:} Active refers to 21.5mg dose of IV SPL026 3: 6-month follow-up out of study

Safety and adverse events

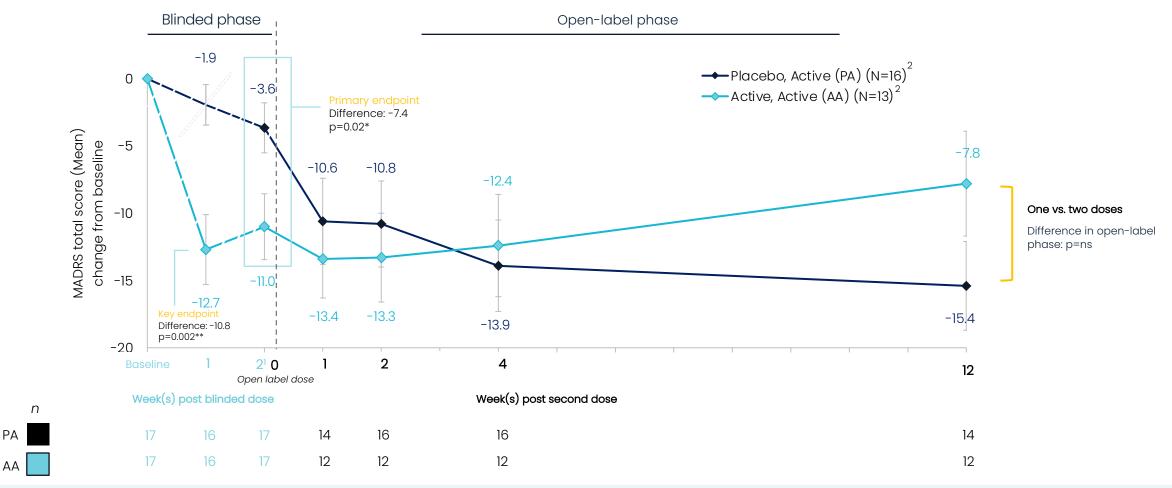
Data suggest *favorable* safety and tolerability profile

- No drug-related SAEs including suicidal ideation or behavior
- 100% of AEs deemed possibly related to treatment were mild to moderate in severity
- The most commonly reported AEs were infusion site pain or reaction, nausea and mild to moderate anxiety
- No clinically significant safety concerns, including no concerns with vital signs, ECG or clinical laboratory findings in any treatment group

Phase IIa: AEs possibly	Blinded pho	Blinded phase (to Day 14)		
related to treatment (n)	Active	Placebo	All subjects	
Infusion site pain or reaction	7	3	17	
Musculoskeletal and connective tissue disorder	1		2	
Nausea	3		6	
Headache	1		2	
Anxiety	2		5	
Insomnia		1	3	
Restlessness	1		2	
Other ¹	6		10	
Total mild and moderate	21	4	47	
Total severe	0	0	0	
Total	21	4	47	



Rapid-onset and durable antidepressant effects of one and two dose regimens of IV SPL026 with supportive therapy



Note: Dashed lines on chart represent blinded phase. Error bars represent Standard Error Mean (SEM); MADRS = Montgomery-Asberg Depression Rating Scale; n = number of data points; N = population number; ns = not significant

a-c: See Appendix – Footnotes and Sources and "Cautionary Notes – Forward-Looking Information", "Risk Factors" and "Treatment Claims"

c: See Appendix – Footnotes and Sources and Cautionary Notes – Forward-Looking information , "Risk Factors" and Treat

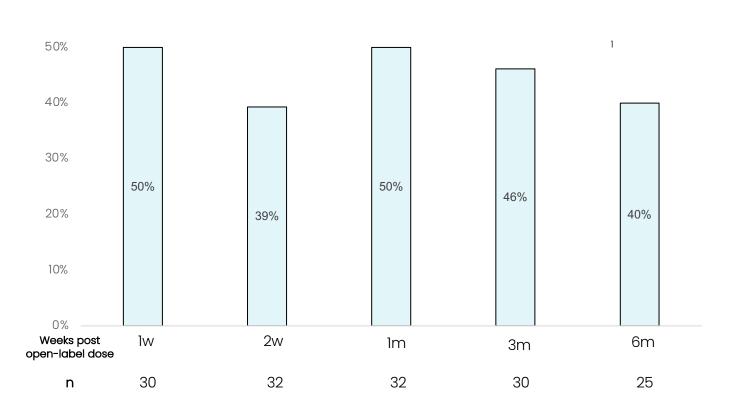
1: Represents Week 2 endpoint of both treatment groups in the blinded phase taken prior to receiving open label dose

^{1:} Represents week 2 enapoint of both treatment groups in the blinded phase taken prior to receiving open label abse 2: N sizes as follows: Blinded phase: placebo arm (N=17) active arm: (N=17). Open-label phase: PA (N=16) and AA (N=13) because 5 participants from blinded phase did not receive open-label dose

Durable remission rates in one and two dose SPL026 regimensa,b,c

Remitters (%) (MADRS score ≤10) - Aggregated¹

60%



Key takeaways

- Outcomes demonstrated in remission data is consistent with response data³ in one and two dose regimens
- Encouraging remission rates demonstrated to at least 6 months
- Among the patients who had achieved remission within three months with SPL026, 64% sustained remission to six months²

Notes: MADRS = Montgomery-Asberg Depression Rating Scale; n = number of datapoints

a-c: See Appendix - Footnotes and Sources and "Cautionary Notes - Forward-Looking Information", "Risk Factors" and "Treatment Claims"

Refers to mean aggregated outcomes of all patients receiving an active dose in the open label phase; and includes four participants excluded from the formal statistical analysis who received a blinded dose of SPL026 but did not receive a second open label dose.

^{2:} Based on data from patients followed up out of study

^{3:} See Corporate Presentation - Small Pharma Phase IIa Topline Results - on website for further details

Analysis of secondary and exploratory measures strengthens primary MADRS efficacy results^{a,b,c,1}

Consistency between patient-reported and independent rater depression scores

Patient self-reported BDI depression scores corroborate MADRS assessments conducted by independent clinical raters. Improvements in BDI mean total score observed across all time points in both one and two dose regimen groups.

Improvements in anxiety scores

Anxiety is often impacted by depression. We observed rapid and statistically significant improvements in anxiety symptoms versus placebo across all time points in the one and two dose regimens, as measured by the STAI-T scale.

Positive impact on patient wellbeing

Wellbeing is also often impacted by depression. We observed a rapid and sustained improvement in patient wellbeing versus placebo across all time points in both the one and two regimen groups, as measured using WEMWBS.



SPL028 a,b,c Deuterated DMT



Profile

- · Novel chemically engineered DMT with deuterium
- Affects rate of drug metabolism in the body
- Anticipated to extend psychedelic experience vs. SPL026

Targeting a DMT-based treatment with a differentiated profile

Anticipated that its distinct pharmacokinetic profile could offer 1) an extended DMT psychedelic experience with the potential for unique clinical benefits and 2) the ability to optimise dosage formulations for various administration routes.

Strong commercial proposition

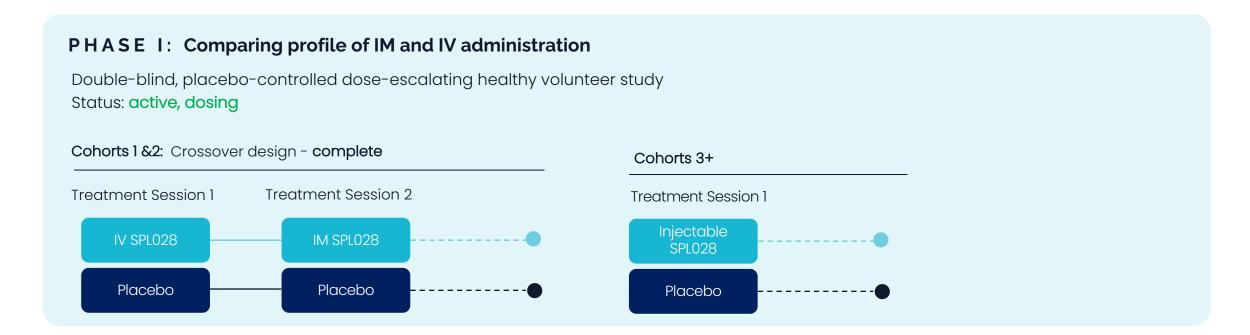
Findings from SPL028 and SPL026 trials to date shows potential for a short in-clinic treatment (~<2.5hour) administered on an episodic "as-required" basis, offering convenience for both patients and physicians.

2nd generation psychedelic with multi-layered IP protection

Maturing IP portfolio surrounding SPL028 and related deuterated compounds including granted Composition of Matter protection in multiple jurisdictions.

17

SPL028 data to date supports target treatment profile a,b,c



→ Key Insights

Non-clinical data

Similar pharmacology to SPL026 when comparing binding profiles against 5-HT receptor subtypes and *in vitro* receptor binding profiles across additional receptors

Preliminary data of ongoing Phase I study (cohorts 1&2)

- ✓ IV SPL028 demonstrates an extended DMT psychedelic experience < 1 hour</p>
- ✓ Well tolerated in subjects dosed to date



Small Pharma Company Highlights^{a,b,c}

Focus on shortduration psychedelics

Advancing portfolio of short-duration psychedelics for mental health disorders that have the potential for rapid-acting and long-lasting relief addressing areas of critical unmet need in mental health.

2

Achieved proof-of-concept for DMT therapy in depression

Demonstrated

favorable safety

profile of a DMT-

based treatment,

placebo-controlled

efficacy and

SPL026, with

Disorder in

randomized,

Phase IIa trial.

3

Clinically differentiated **DMT** candidates

4

Robust multi-layered IP portfolio

5

Cash runway to deliver on kev milestones

Clinical findings to date supports potential for differentiated DMTbased drugs that could offer distinct supportive therapy clinical and in Major Depressive commercial benefits

Maturing and expanding IP portfolio with 23 patents granted and 90+ applications pending.1

Cash position anticipated to allow for completion of key value-based milestones in 2023/24.

19

Thank you.

References

GENERAL

- a. Certain statements regarding tryptamine-based treatments have not been evaluated by the U.K. Medicines and Healthcare products Regulatory Agency, the U.S. Food and Drug Administration, Health Canada, or other similar regulatory authorities, nor has the efficacy of DMT-based treatments been confirmed by approved research. There is no assurance that tryptamine can be used to diagnose, treat, cure or prevent any disease or condition and robust scientific research and clinical trials are needed.
- b. Forward-looking statements are subject to various risks and assumptions. See "Cautionary Notes" on page 2 of this presentation.
- c. Subject to receipt of all necessary regulatory approvals from all applicable governmental authorities. There are multiple risk factors regarding the ability to successfully commercially scale and develop DMT-based treatments and a portfolio of DMT analogues

SLIDE 3

- 1. WHO (2021), Depression factsheet
- Rush AJ et al. "Acute and longer-term outcomes in depressed outpatient requiring one or several treatment steps: A STAR*D report". The American Journal of Psychiatry. 2006. 163(11):1905-1917
- 3. Sansone RA, Sansone LA. Antidepressant adherence: are patients taking their medications?
- 4. Davis, A. et al. (2021) Effects of Psilocybin-Assisted Psychotherapy on Major Depressive Disorder
- Holze, F., Caluori, T.V., Vizeli, P. et al. Safety pharmacology of acute LSD administration in healthy subjects. Psychopharmacology (2021)
- 6. Refer to Slides 11-16 for further information on the SPL026 program
- 7. Refer to Slides 17-19 for further information on the SPL028 program

SLIDE 4

- WHO (2021), Depression factsheet
- 2. American Association of Suicidology, 2014
- 3. Arnaud et al (2021) The Increasing Economic Burden with Additional Steps of Pharmacotherapy in Major Depressive Disorder
- 4. Rush AJ et al. "Acute and longer-term outcomes in depressed outpatient requiring one or several treatment steps: A STAR*D report". The American Journal of Psychiatry. 2006. 163(11):1905-1917

Definitions as defined in the STAR*D trial:

Efficacy rate (Remission): Quick Inventory of Depressive Symptomatology, QIDS-SR-16, scale was administered at each clinic visit, and remission was measured as a score of ≤5

Intolerance: Patients who failed to complete at least 4 weeks of treatment

Relapse: QIDS-SR 16 score ≥11 (corresponding to a Hamilton Depression Rating Scale, HDRS-17 ≥14)

