Supernus Pharmaceuticals



Jefferies Healthcare Conference

June 2022



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Proven Execution in CNS & ADHD

30+ Years of CNS experience including Four Products in ADHD



2005 - Present



















SPN-830

SPN-820

SPN-817

SPN-840

SPN-443

SPN-446

SPN-448

SPN-396



1997 - 2005









For several years, and prior to becoming independent in 2005, Supernus operated as Shire Laboratories, Inc., a division of Shire. SPN-830, SPN-820, SPN-8417, SPN-840, SPN-443, SPN-446, SPN-448, and SPN-396 are product candidates in various stages of development. All trademarks are the property of their respective owners



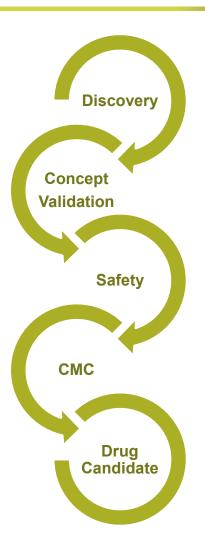
Significant Experience & Capabilities in Drug Development

Discovery Platform

Design and synthesis of new compounds based on structure, function and disease pathways

Toxicology

Validated drug candidates

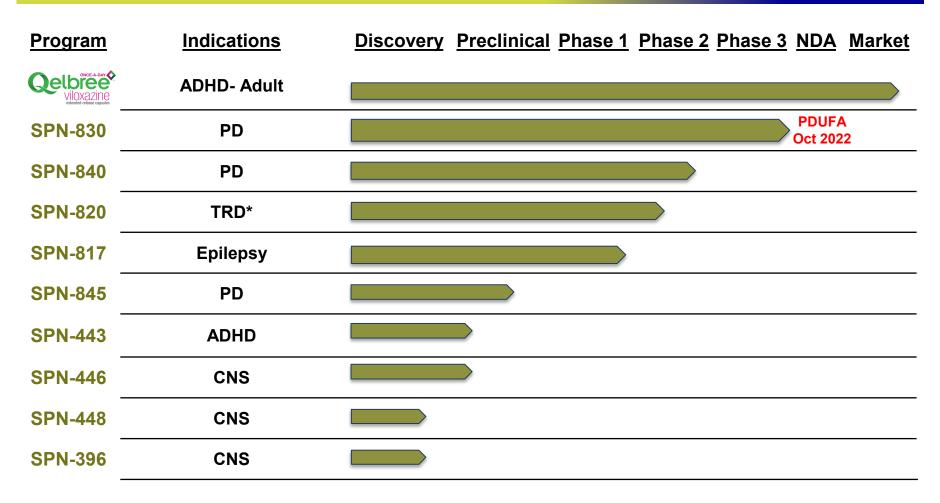


In vitro, PK, preclinical proof of concept, and stability studies

In-house CMC/drug delivery expertise & GMP manufacturing



Robust CNS Pipeline to Drive Long-term Growth



^{*}TRD = Treatment Resistant Depression PD = Parkinson's Disease





Novel Non-Stimulant ADHD Product

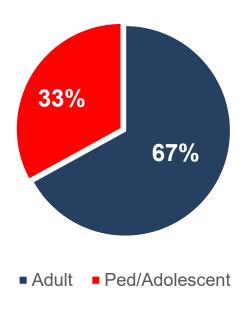
- Launched in May 2021 for patients 6 to 17 years of age
 - Sales force of approximately 195 sales representatives
 - Clinical feedback from the market in line with Phase III clinical data demonstrating a well-differentiated profile
- Launched in May 2022 for adult patients
- IP with expirations from 2029-2033
- ADHD market is significant with 80 million prescriptions/year¹



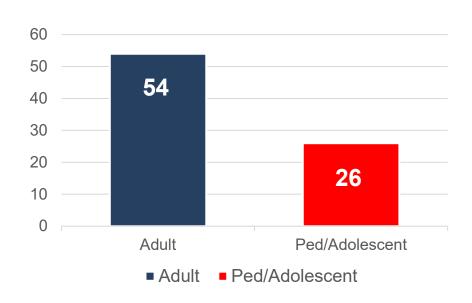
ADHD Adult Opportunity: Significant Market Segment

2021 Total ADHD U.S Market - 80 Million Prescriptions

% of Market



2021 Prescriptions (millions)

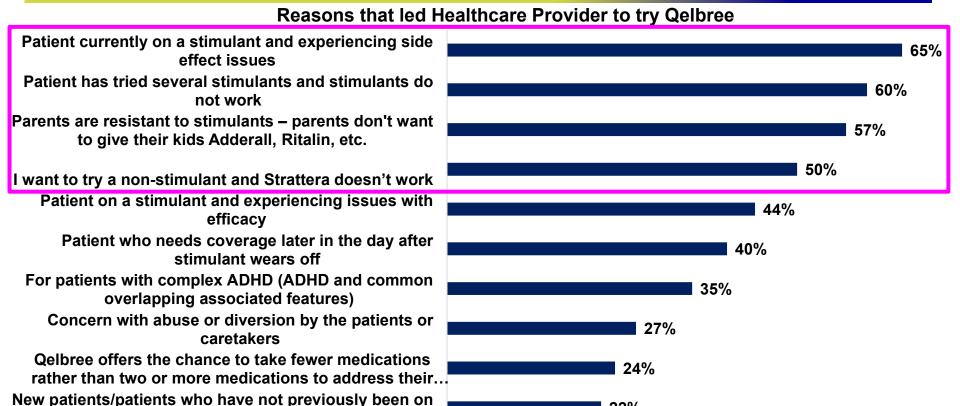


Source: IQVIA



Top Reasons to Try Qelbree®

Patients Having Issues With Stimulants & Looking For a Non-Stimulant That Works



22%

19%

Source: Internal market research among Qelbree prescribers – n=104.

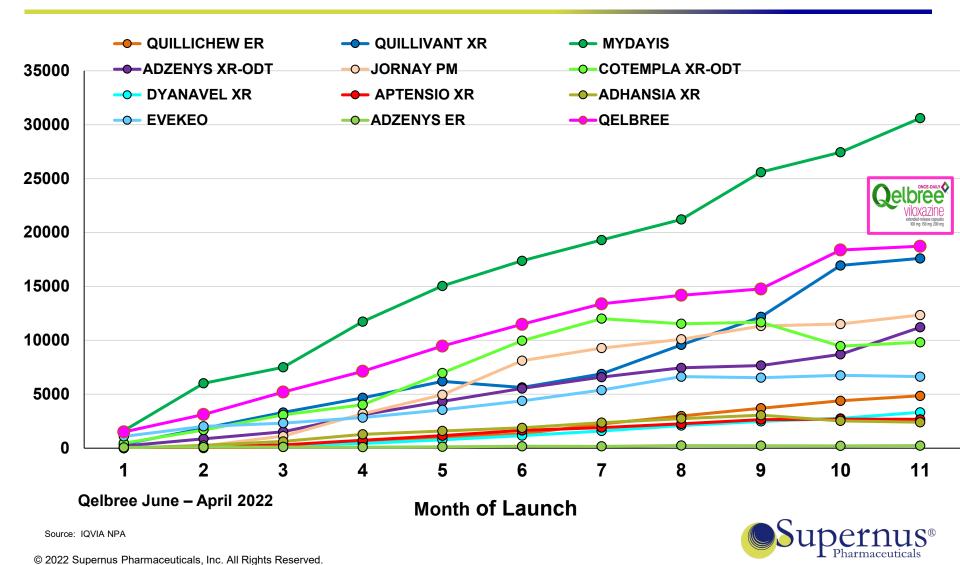
Q28. Thinking about the patients you have put on Qelbree, what medications were they on/what issues were they having that led you to try Qelbree?



medication

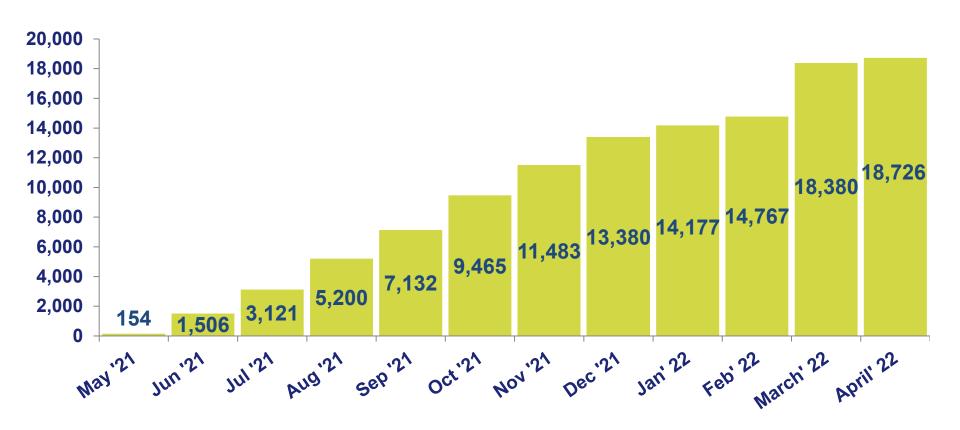
Patient needed something once a day

ADHD Launch-Aligned Monthly Prescriptions





Monthly Prescriptions since Launch



Source: IQVIA NPA



Major Presence in Parkinson's Disease (PD)

1 Million U.S. PD Patients - Market Expected to Grow to \$6.2B by 2026 (1)





Significantly Decreased Dyskinesia and OFF Time, Thereby Significantly Increasing Good ON Time

GOCOVRI achieved reductions in dyskinesia & OFF episodes without having to adjust levodopa dose

Placebo-adjusted, pooled results from pivotal trials*

Primary endpoint

L27%
DECREASE IN DYSKINESIA

10.1-point reduction in UDysRS score

 $(-17.7 \text{ GOCOVRI vs. } -7.6 \text{ placebo})^{(1)(2)\dagger}$

- (1) Elmer LW, CNS Drugs. 2018.
- (2) Data on file. Adamas Pharma LLC, Emeryville, CA.

* Pooled results from 2 independent positive, pivotal, Phase 3, randomized, placebo-controlled trials (Study 1 and Study 2) in PD patients on levodopa. Study 1, a 24-week study, was conducted in 121 PD patients with dyskinesia (GOCOVRI [n = 63], placebo [n = 58]). Study 2, a 12-week study, was conducted in 75 PD patients with dyskinesia (GOCOVRI [n = 37], placebo [n = 38]).

 \uparrow In Study 1, GOCOVRI reduced the UDysRS total score by 15.9 points (vs 8.0 with placebo) (P = 0.0009), decreased OFF time by 0.6 hours (vs an increase of 0.3 hours with placebo) (P = 0.0171), and increased GOOD ON time by 3.6 hours (vs 0.8 hours with placebo) (P < 0.0001) from baseline. In Study 2, GOCOVRI reduced the UDysRS total score by 20.7 points (vs 6.3 with placebo) (P < 0.0001), decreased OFF time by 0.5 hours (vs an increase of 0.6 hours with placebo)

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Secondary endpoints

136%
DECREASE IN OFF TIME

1-hour decrease

 $(-0.6 \text{ GOCOVRI vs. } 0.4 \text{ placebo})^{(1)(2)\dagger}$

129%
INCREASE IN GOOD ON TIME

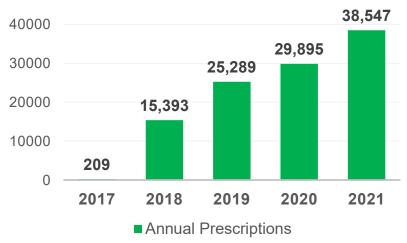
2.4-hour increase

(3.8 GOCOVRI vs.1.4 placebo) (1)(2)†



A Key Growth Driver

- 2021 Net Sales : \$87.6M, 23% growth vs. 2020
 - Supernus recorded \$9.8M in 2021 net sales (acquisition closed on Nov 24, 2021)
- For the treatment of dyskinesia in patients with PD receiving levodopa-based therapy, with or without concomitant dopaminergic medications
- Adjunctive treatment to levodopa/carbidopa in patients with PD experiencing "off" episodes



Source: Company Records





Significant Target Patient Population

Over 50% of people with PD experience OFF episodes, dyskinesia or both within 5 years, and up to 100% after 10 years (1)(2)

GOCOVRI potential addressable U.S. patient population

400,000 to 500,000 patients(3)

- (1) Kim H-J, et al., Mov Disord, 2020.
- (2) Mizuno Y et al., Journal of Neural Transmission, 2018
- (3) Estimated based on market research.

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1,000,000

PD PATIENTS DIAGNOSED IN U.S.

800,000

DIAGNOSED AND TREATED PATIENTS

700,000

LEVODOPA-TREATED PATIENTS





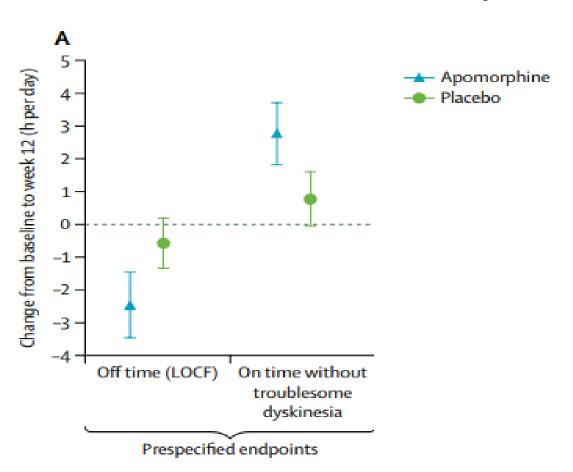
Novel Apomorphine Subcutaneous Injection Device

- Non-invasive dopaminergic stimulation therapy for continuous treatment of ON-OFF episodes in PD
- Potentially first non-surgical continuous dopaminergic stimulation device
- Currently available options
 - Gastro-intestinal surgically implanted levodopa/carbidopa infusion
 - Deep brain stimulation
- Could be eligible for Orphan Drug Designation and 7-year exclusivity
- NDA filing accepted by FDA with early October 2022 PDUFA



Novel Apomorphine Subcutaneous Injection Device

TOLEDO Phase III Study Results



Primary Endpoint

SPN-830 demonstrated a
2.47 hours per day
reduction in OFF time
compared to placebo (0.58);
p= 0.0025

Regina Katzenschlager et al, The Lancet Neurology. 2018; Vol 17(9):749-759

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Novel MOA for Treatment-Resistant Depression (TRD)

- First-in-class selective brain mTORC1 activator
 - Directly activates mTORC1 enhancing synaptic activity and cellular metabolism in the brain
- Early efficacy signal on HAMD-6 scale in TRD patients
 - Rapid onset of action (signal at 2 hours)
 - Meaningful effect sizes (>0.4 through 3 days on 1 dose)
- Multiple ascending dose (MAD) study demonstrated:
 - drug penetration and target engagement
 - favorable safety/tolerability profile across broad range of doses
- Initiated Phase II clinical trial in TRD
- Significant market opportunity
 - Major depressive disorder (MDD) affects approximately 17.3M U.S. adults¹
 - ~30% of MDD patients are treatment resistant²



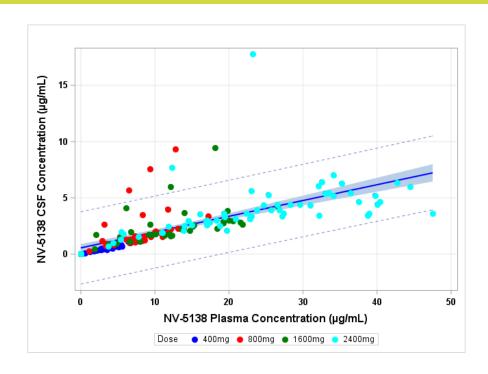
¹⁻ National Institute of Mental Health, 2017 National Survey on Drug Use and health
2- Rush AJ Et al.(2006) Acute and Longer-Term Outcomes in Depressed Outpatients Requiring One or Several Treatment Steps: A STAR*D Report. The American Journal of Psychiatry,163(11), 1905-1917

Multiple Ascending Dose Study

- Healthy subjects, placebo-controlled pharmacokinetic study
 - 5 Cohorts: 400 mg, 800 mg, 1600 mg, 2400 mg and 3000 mg
- Plasma & cerebrospinal fluid (CSF) drug concentration
 - 400 mg 2400 mg dose levels
 - Single dose, at Day 1
 - Multiple doses, at Day 7
- Metabolomic biomarkers of mTORC1 activation
 - Concentrations measured in CSF for 400 mg 2400 mg dose levels
 - N-acetylmethionine
 - N-formylmethionine
 - Orotic acid



SPN-820 CSF Concentration vs. Plasma Concentration

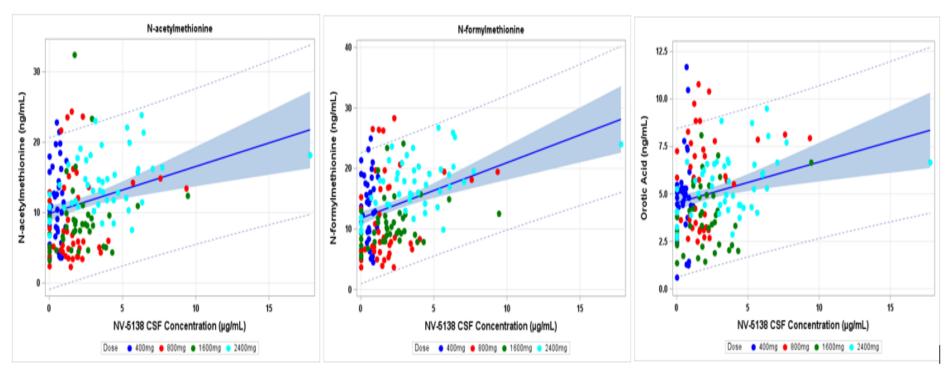


SPN-820/NV-5138 CSF concentrations significantly increase with the increase in plasma concentrations (p < 0.0001)

Dots represent the observed data, the solid line represent the model predicted curve, the shaded area represent the 95% confidence interval on the predicted curve, and the dotted lines delimit the 95% prediction interval.



Biomarker Concentrations vs. SPN-820 CSF Concentrations



Dots represent the observed data, the solid line represent the model predicted curve, the shaded area represent the 95% confidence interval on the predicted curve, and the dotted lines delimit the 95% prediction interval.

Biomarker concentrations significantly increase with the increase of SPN-820/NV-5138 CSF concentrations



Financial Summary and Guidance

1Q 2022 Financial Results¹

- Total revenues of \$152.5 million, up 16% over 1Q 2021
- Net earnings and diluted earnings per share (GAAP) of \$25.6 million and \$0.43, respectively
- Operating income (GAAP) of \$2.0 million; operating income (non-GAAP) of \$28.0 million
- Cash, cash equivalents, and investments at \$437.5 million as of March 31, 2022



¹ Reported on May 9, 2022, including reconciliation of GAAP vs Non-GAAP

Financial Summary and Guidance

Full Year 2022 Financial Guidance¹

	(\$ millions)
Total Revenues	\$640 - \$680
Combined R&D and SG&A expenses	\$460 - \$490
Operating Earnings - GAAP	\$20 - \$40
Adjustments:	
Amortization of intangible assets	\$80 - \$85
Share-based compensation	\$20 - \$25
Contingent consideration	\$8 - \$12
Depreciation	\$2 - \$3
Operating Earnings - non-GAAP	\$130 - \$165



¹ Guidance as confirmed on May 9, 2022

Positioned For Long-Term Growth



Diversified CNS Portfolio

Qelbree®, Oxtellar XR®, Trokendi XR®, APOKYN®, GOCOVRI®, XADAGO®, MYOBLOC®

Innovative Pipeline in CNS

Qelbree® ADHD (Adult)

 SPN-830
 PD

 SPN-820
 TRD

 SPN-840/845
 PD

SPN-817 Epilepsy

SPN-443 ADHD

SPN-446/448/396 CNS

