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SPARC timeline



Built a robust R&D engine over 15 years

Integrated translation advancing standard of care in difficult-to-treat diseases

- Incubated out of Sun Pharma, India's largest pharmaceutical company
- Spun out in 2007 to an independent listed entity
- Focus on exploratory programs and capability development

- Initial focus on Medicinal Chemistry and Formulations
- Success with Lipodox
- Multiple delivery systems innovations
- Substantial investments in discovery & translational development

- US FDA Approvals of Xelpros and Elepsia
- NCEs enter clinical Development
- Focus on strategic partnering to access early science

- Growing NCE pipeline
- 3 more NDA submissions
- Forays into ADCs & Immunofusions
- Poised to grow its multimodality platform in CNS, Oncology & Immunology

2007-2010

2011-2014

2015-2019

2020 & beyond



Investment highlights





4 Clinical Stage Programs Targeting Areas of High Unmet Need

Targeting large addressable patient populations with USD 20Bn+ combined peak sales potential in 6 indications within Oncology, Neurology, and Immunology



Discovery & Development Across Validated & Novel Biology in Order to Balance the Risk

- Multi-modal portfolio covering small and large molecules and conjugated entities
- 10+ preclinical programs including an ADC program expected to enter the clinic in 2023



Proven High Quality R&D Organization with Capital-Efficient Global Operations

- 350+ scientists across 4 research centers including USD 400Mn invested to date
- 2 USFDA approvals for internally developed assets
- 3 NDAs targeted for submission in 2022



Highly Flexible Model to Maximize Shareholder Value

- Partnerships to maximize large commercial potential and provide non-dilutive capital
- Maximize multi-TA opportunity and preserve optionality for spin-offs



Experienced Management Team and Globally Recognized Scientific Advisory Board









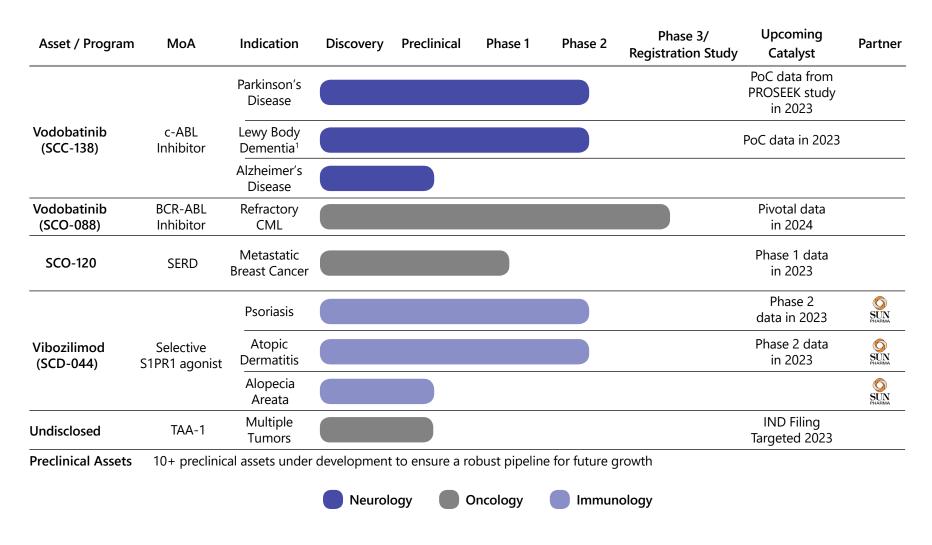






Pipeline overview & key milestones





^{1.} Investigator Initiated Study | MoA = Mechanism of Action | PoC = Proof of Concept | CML = Chronic Myeloid Leukemia | SERD = Selective Estrogen Receptor Degrader S1PR1 = Sphingosine-1-Phosphate Receptor 1 | IND = Investigational New Drug | TAA-1 = Tumor Associated Antigen-1



Vodobatinib for Neurodegenerative Diseases (SCC-138)

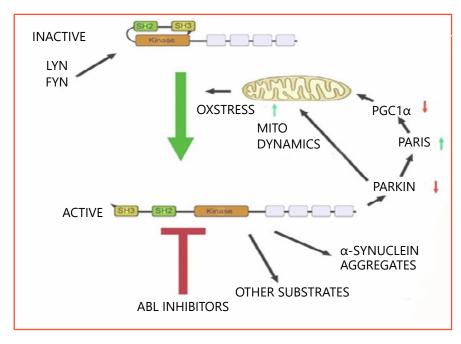
A potential first-in-class disease modifying therapy

Vodobatinib for neurodegenerative diseases



Optimal agent to test the c-Abl hypothesis

Mechanism of Action of c-Abl inhibition



Reduces neuronal toxicity caused by the aggregated neurotoxic proteins

- Vodobatinb is a potential first-in-class c-Abl inhibitor for Parkinson's disease
- Augments autophagic flux and prevents inactivation of Parkin-mediated mitochondrial quality control
- Reduces α-synuclein inclusions
- Sub-nanomolar potency against human c-Abl
- Very limited off-target activity, leading to improved safety profile
- Robust brain penetration (Brain/ Plasma levels around 0.9)

Selective Abl inhibition

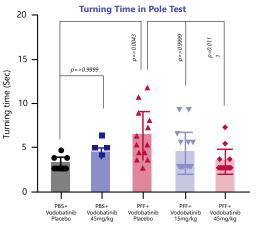
Family	Kinases	IC ₅₀ (nm)
Abl	Abl (Abl-1)	0.9 0.8
	Arg (Abl-2)	0.0
SFK	Src	90.0
	Fyn	18.0
	Hck	54.0
	Lck	17.0
	Lyn	18.0
	Yes	28.0
	PTK5	3.0

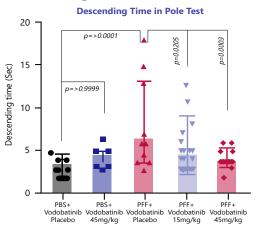
Behavioral assessments in the PFFinduced mouse model



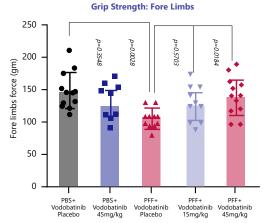
- In the MPTP¹ mouse model,
 Vodobatinib prevents neuronal degeneration in substantia nigra
- In the PFF² induced mouse model, vodobatinib shows target engagement, reduction in Serine 129 phosphorylation of α-Synuclein, preservation of dopaminergic neurons and clinical improvement in motor and cognitive functions
- In the AAV³ driven rat A53T α-synuclein model, vodobatinib shows neuroprotection

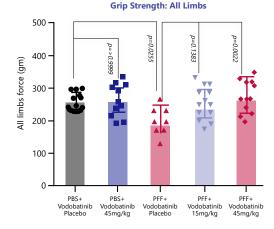
Vodobatinib at 45 mg/kg improves PFF-induced movement disorderrelated deficits in Turning Time and Descending Time in the Pole test





Vodobatinib treatment improves PFF-induced deficits in Grip Strength

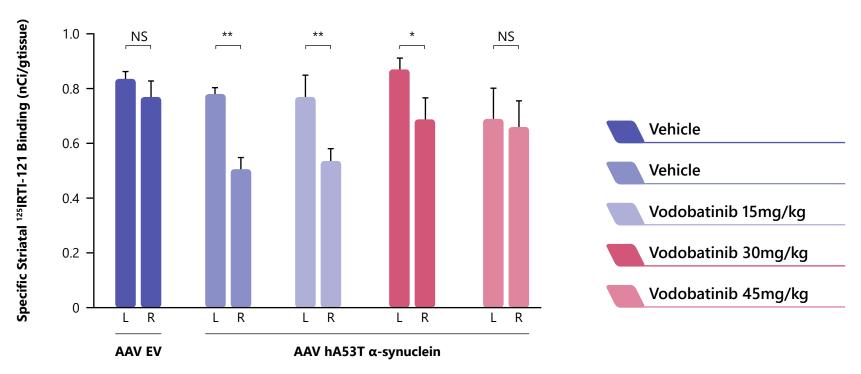




^{1.} Data generated in-house | 2. Study conducted at the Ted Dawson Lab, Johns Hopkins University | 3. Study conducted by Atuka Inc. Unpublished data; not to be replicated or shared | PBS = Phosphate-buffered saline | PFF = Preformed fibril

Vodobatinib protects dopaminergic neurons in the AAV mutant α -synuclein (hA53T) rat model -dopamine transporter expression





NS: p>0.05; *p<0.05; *p<0.001 versus the un-operated (contralateral) hemisphere. Two-way ANOVA with Fisher's LSD post-hoc test

- Vodobatinib treatment protects against dopaminergic neuronal loss measured by radiolabeled ¹²⁵I labeled RTI-121 binding in the striatum
 - Comparison of un-operated left hemisphere (L) and operated right hemisphere (R, injected with & expressing the AAV) shows that 45 mg/kg doses provides protection of dopaminergic neurons

Vodobatinib met the brain exposure targets in early clinical studies



Summary of completed toxicology, safety pharmacology and clinical studies

Preclinical toxicology update

- Acute tox in mouse and rat by oral route, and in rat by ip route
- Repeat dose oral tox in rat (upto 6 months) and beagle dog (upto 9 months)
- Genotoxicity (In vitro Ames' Test and In vivo mouse micronucleus study)
- Repro toxicity
- Safety Pharmacology, including CVS safety

Clinical summary

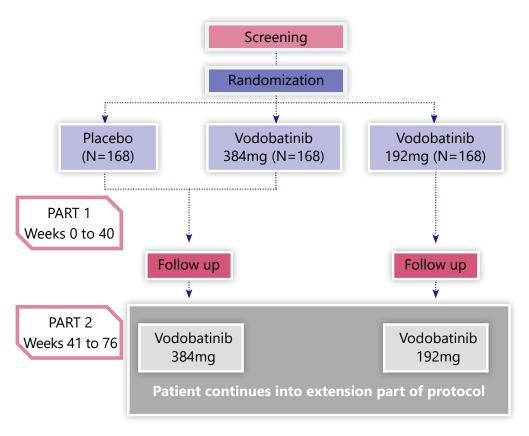
- Phase 1 completed in healthy subjects, PD subjects up to 384mg
 - Overall well tolerated
 - PK suggests adequate brain penetration over 24 hours

Trial	Population	Status	Safety findings
Phase 1 MAD	PD any stage	cohorts of 8 subjects each on 14 days of Vodobatinib or placebo capsules (6:2 randomization) 6, 12, 24, 48, 96, 192, 384mg	Well tolerated
Phase 1	Healthy men	48, 192mg, 384mg x7 days with 24 hours of CSF sampling on day 7. Study complete	Mild AEs
Phase 1 Crossover study	18 Healthy subjects per cohort	192mg powder vs 192 mg capsule 384mg powder vs 192mg capsule 384mg powder fed vs fasting	No significant concerns

Vodobatinib for Parkinson's Disease



Recruitment on track to achieve Phase 2/PROSEEK enrollment target in 2022



PROSEEK

- 84 sites across US, Europe and India functional; recruitment ongoing to complete enrollment in 2022
- Over 40% patients randomized (N=218)
- Phase 2 readout expected in 2023

Primary outcome

 Change in MDS-UPDRS Part 2 + Part 3 from baseline to end of treatment

Secondary outcomes:

- Time to start of symptomatic medication
- CGIS clinician global impression of severity
- PK/PD correlations

Exploratory outcomes:

- DaT SPECT at beginning (in all subjects for eligibility) and end
- Skin biopsy for synuclein deposition at Baseline and Week 36
- Smartphone-based measure of motor performance
- Exploratory CSF markers

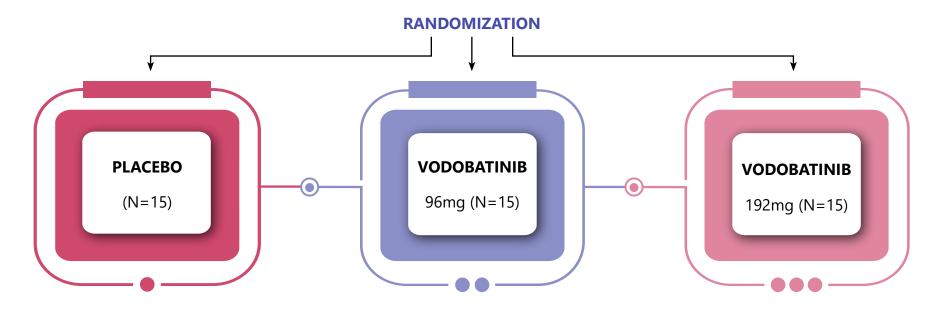
Data cut-off date: 26th Nov 2021 | A Phase 2 Study In Early Parkinson's Disease Patients Evaluating The Safety And Efficacy Of Abl Tyrosine Kinase Inhibition Using K0706 K0706 = Vodobatinib PD (SCC-138) | NCT03655236

Opportunities beyond Parkinson's Disease



Dementia with Lewy Bodies offers an immediate next opportunity

- ODLB is a neurodegenerative condition with progressive cognitive impairment, hallucinations and parkinsonism
 - Estimated to affect about 1.4 million people in the USA*
 - 2nd most common cause of dementia in the elderly
- Strong overlap with Parkinson's Disease
- Synucleinopathies with Lewy Bodies seen on autopsy. Pathophysiology similar to PD suggesting potential efficacy in DLB
- Investigator-initiated trial in collaboration with Georgetown University, Washington on-going in subjects with DLB





Vibozilimod (SCD-044) -A Selective S1PR1 Agonist

A safer alternative to JAK inhibitors

Vibozilimod (SCD-044)



An opportunity to improve oral standard of care in dermatology

Vibozilimod is a Best-in-Class S1PR1 modulator with excellent safety

S1PR1 Modulator Landscape

- Fingolimod is the First-in-Class S1PR agonist approved, but being a non-selective modulator, is associated with serious cardiac side-effects
- Multiple S1PR1 modulators are approved (siponimod and ozanimod) for non-dermatology indications; vibozilimod has opportunity to lead the field in dermatology
- Recent safety concerns related to JAK inhibitors (including topical/locally delivered agents) increase the significance of S1PR1 agonists as a 'class alternative' in several autoimmune disorders, particularly in dermatology

Vibozilimod (SCD-044)

- Developed in collaboration with a French biotech company,
 Bioprojet SPARC in-licensed Bioprojet's share of IP in 2019
- Highly-selective for S1PR1 over S1PR2 and S1PR3, which can be associated with serious side effects
- Established preclinical and early clinical validation
- Currently targeting atopic dermatitis, psoriasis and other autoimmune disorders
- Potential synergy with other mechanisms in IBD like IL-23 blockade

S1PR1 agonists	EC ₅₀		
STERT agonists	S1PR1	S1PR3	S1PR5
Vibozilimod ¹	0.2	>10,000	9
Fingolimod ¹	0.4	7.7	2.2
Ozanimod ¹	1.9	>10,000	3.5
Ponesimod ¹	~1	NA	10.7
Etrasimod ¹	1.5	~1000	0.7

Vibozilimod licensed to Sun Pharma with around ~50% economics retention

Vibozilimod (SCD-044)



Pharmacodynamics and safety established in Phase 1 study

Multi-part Phase 1 study completed in healthy volunteers

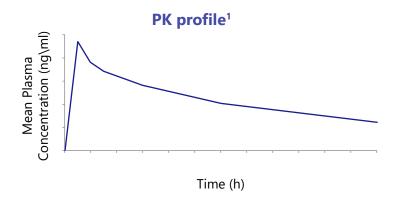
Single Ascending Dose

- Six dose levels in males and one dose level in females
- ~55% lymphocyte count decrease following 1 mg dose

Multiple Ascending Dose

- Four dose levels including two dose up-titration schemes in males and one dose up-titration scheme in females
- ~60% lymphocyte count reduction observed at 1 mg dose with asymptomatic bradycardia
- Reduction in lymphocyte count confirms potential efficacy of vibozilimod

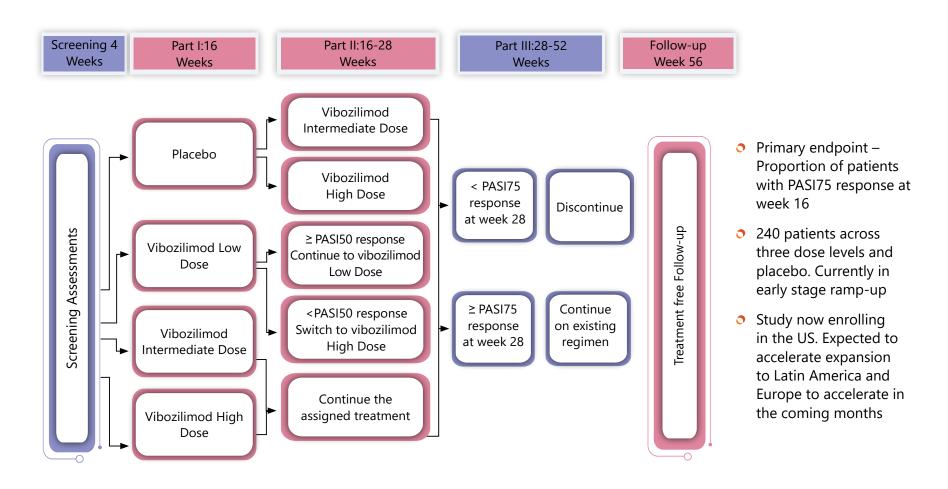
Change from Baseline Tymphocyte count reduction Tymphocyte count reduction Tymphocyte count reduction Tymphocyte count reduction Tymphocyte count reduction



Vibozilimod (SCD-044) for psoriasis

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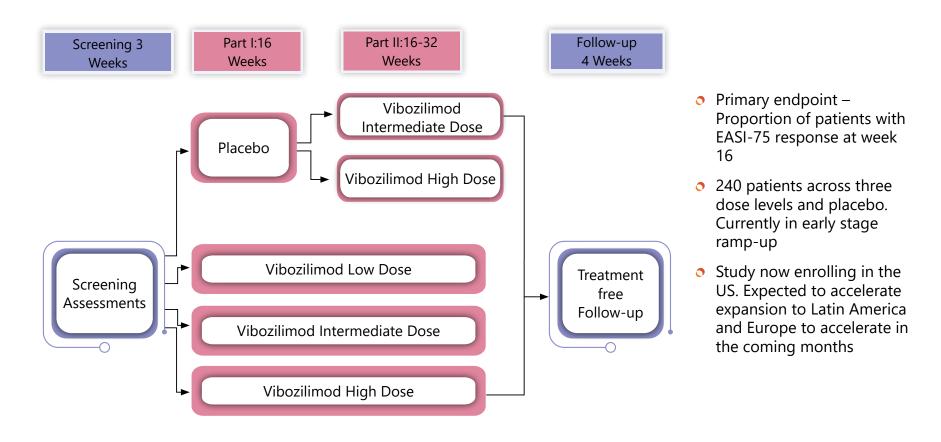
Clinical proof-of-concept by 2023



Vibozilimod (SCD-044) for atopic dermatitis



Clinical proof-of-concept by 2023





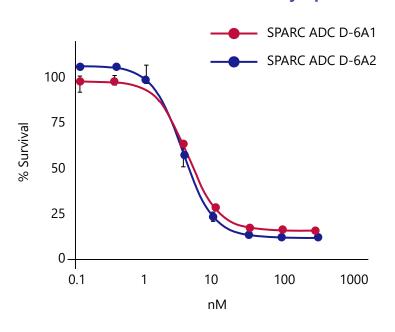
Anti TAA-1 Asset

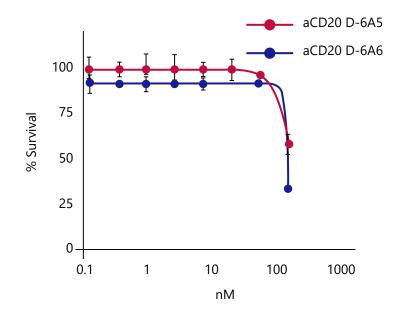
Targeting an antigen expressed in a wide spectrum of tumors

SPARC ADC binds and exerts cytotoxicity against target-expressing cells



Cytopathic assay in a pancreatic cancer cell line





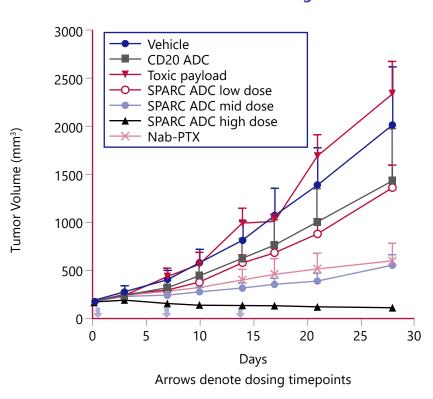
- ADC against a novel tumor associated antigen as a target
- Evidence of potent cytotoxicity of SPARC ADC against TAA-1 over-expressing pancreatic carcinoma cell line
- 100-fold greater potency over a nonbinding ADC of the same payload targeted to CD20

Antitumor efficacy of SPARC ADC

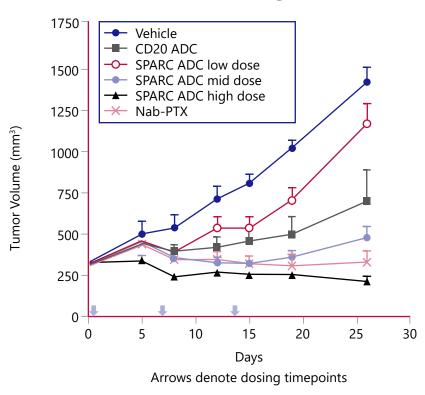


Efficacy established in multiple xenograft models

Pancreatic carcinoma xenograft



Ovarian carcinoma xenograft



- Dose-dependent growth inhibition of xenografts of pancreatic and ovarian carcinomas using SPARC ADC
- Control nonbinding anti-CD20 ADC as well as unconjugated cytotoxic agent were ineffective

SPARC ADC: next steps



- Advance anti TAA-1 ADC through preclinical development with IND submission in 2023
- Explore additional tumor-targeting specificities for creation of drug conjugates
- In light of the broad expression of TAA-1 in cancer, create and preclinically evaluate a series of additional immunefusions anchored on TAA-1 targeting
 - TAA-1 targeted T-cell engager (TCE)
 - Bispecific TAA-1 targeted immune-fusion with anti-angiogenesis activity of TCEs
 - Bifunctional TAA –1 targeted immunocytokine(s) to enhance antitumor activity
 - TAA-1 targeted nanoparticles for preferential tumor-focused delivery of other targeted agents

Potential for multiple biologic product INDs in the next five years



Vodobatinib in CML (SCO-088)

A safer, last-line option for heavily pre-treated patients

Vodobatinib for CML (SCO-088)

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Promising Last Line Therapy









- CML is caused by a translocation of the abl gene that results in formation of Philadelphia Chromosome
- Prior to the discovery of BCR-ABL inhibitors, CML was a fatal disease with an 8-year survival rate of ~6%
- Tyrosine kinase inhibitors have changed the prognosis of CML, but patients eventually can become resistant to drugs
- Annual incidence of CML is likely to increase at a rate of 1–2 cases per 100,000 adults, est. 8,000 people in US in 2020¹

- Branded 2nd and 3rd generation TKIs retain high commercial value due to refractory nature of CML, despite genericization of 1st generation TKI
- Large market opportunity US drug sales of the CML TKIs over \$3Bn²
- Our Unmet need for a potent and safe drug in patients with ≥ 3 lines of failure including failure of Ponatinib, given
 - Almost half of patients will have recurrence within 5 years of initial therapy
 - One-third of 2nd line patients and est. 40% of 3rd line patients are refractory or relapse within a year of initiation of that line of therapy

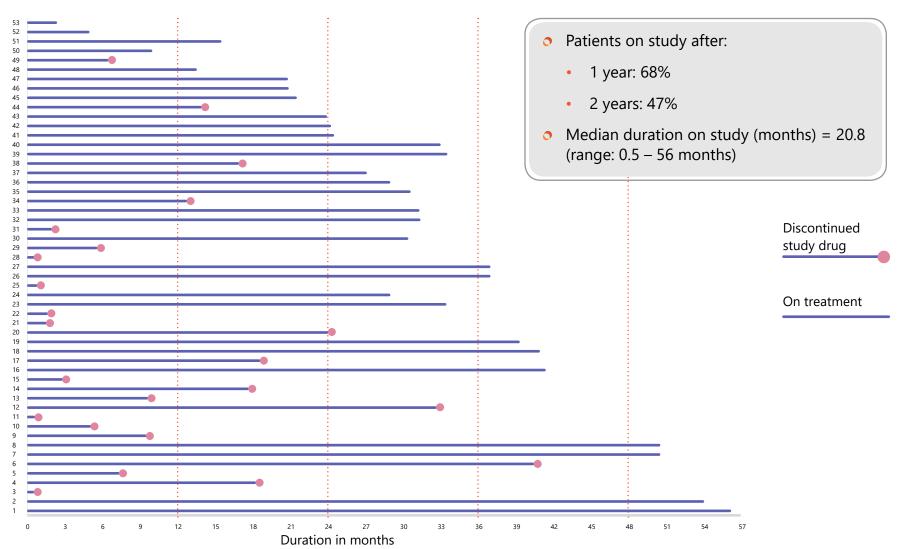
- Targeting patients who are refractory and/or intolerant to other TKIs
- Well tolerated with significant coverage of the mutational field
- Has shown promising activity in clinical trials
- Orphan Drug
 Designation and
 Accelerated Approval pathway agreed with USFDA

- Phase 1 completed in CML subjects
- Favorable safety and tolerability
- Registration study underway. Planned US NDA filing in 2024

Vodobatinib for CML (SCO-088)

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Durable long-term responses seen across cohorts



Vodobatinib for CML (SCO-088)



Clinical Development Plan

Pivotal (Part C) study ongoing

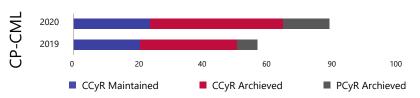
Clinical Development Plan

Part A	Single Ascending Dose study (SAD) in volunteers	⊘
Part B	Multiple Ascending Dose study (MAD) in patients	\odot
Part C	Pivotal efficacy study in refractory and/or intolerant patients to 3 prior TKIs	

- Orphan Drug Designation approved by USFDA and EMA
 - Market exclusivity in addition to IP coverage
 - User fee waiver
- EOP1 discussion completed; agreement with USFDA reached on accelerated approval pathway based on Part C (pivotal study)

Efficacy

Cytogenetic Response (% patients with MaCyR)

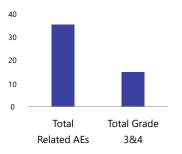


- Major Cytogenetic response in 67% of the enrolled subjects
- Major Cytogenetic response in 54% of the enrolled subjects that meet pivotal study criteria

Safety and Tolerability

 Generally well tolerated with slight excess of GI and hematological AEs





Planned US NDA filing in 2024

EOP1 = End of Phase 1 | MaCyR = Major Cytogenetic Response | CP = Chronic Phase | CCyR = Complete Cytogenetic Response | PCyR = Partial Cytogenetic Response | AE = Adverse Event | GI = Gastro Intestinal SAD = Single Ascending Dose | MAD = Multiple Ascending Dose.



SCO-120 for HR+/HER2-MBC

Potent oral SERD with preferential brain penetration

Oral SERD for Breast Cancer (SCO-120)











- Breast cancer is the second most common cancer diagnosed in women in the United States¹
- Annual incidence of ~2 million patients across the world¹
- ~70% of the breast cancer is HR+/HER2-1
- Hormonal therapy is SoC for ~70% of HR+/HER2metastatic breast cancer patients 1. ERα mutations develop in 20–50% of patients with metastatic disease
- Treated mostly with SERMs, 20–50% patients experience mutations or become resistant
- SERD can break down receptors and prevent cells from dividing. IM Fulvestrant is the only approved SERD but it is poorly active against mutations at therapeutic dose

- SCO-120 is a novel orally-active SERD for the treatment of HR+/HER2- breast cancer
- Active in vitro (nM to sub nM potency) and in vivo in xenograft models against WT ERa and its mutants Y537S and D538G
- In vitro and in vivo studies have shown potential for combination with CDK4/6 inhibitors (palbociclib) in both the WT ERa and the mutation setting
- Favorable Tox profile; No adverse effects seen in battery of in vivo safety pharmacology studies of central nervous system, cardiovascular system, and respiratory system

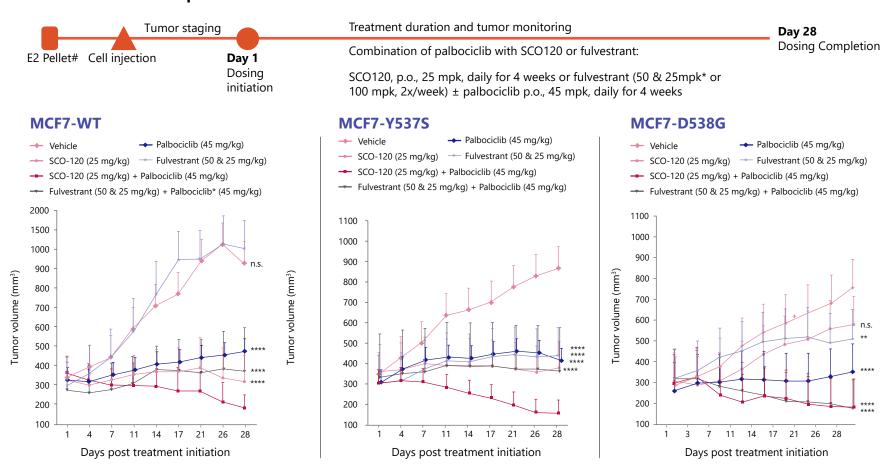
- US IND filed in Jan 2020
- SAD and MAD in healthy volunteers ongoing
- 50 1200 mg cohorts completed. Generally safe and well tolerated, no significant AEs

^{1.} CancerMPact® Treatment Architecture U.S., Breast Cancer | HR = Hormone Receptor | HER2 = Human Epidermal Growth Factor Receptor 2 | ERα = Estrogen Receptor α | SOC = Standard of Care | IM = Intramuscular SERD = Selective Estrogen Receptor Degrader | AE = Adverse Event | SERM=Selective Estrogen Receptor Modulator | MAD=Multiple Ascending Dose

In vivo efficacy of SCO-120 in combination with palbociclib



Promising activity against resistant mutants alone and in combination with palbociclib

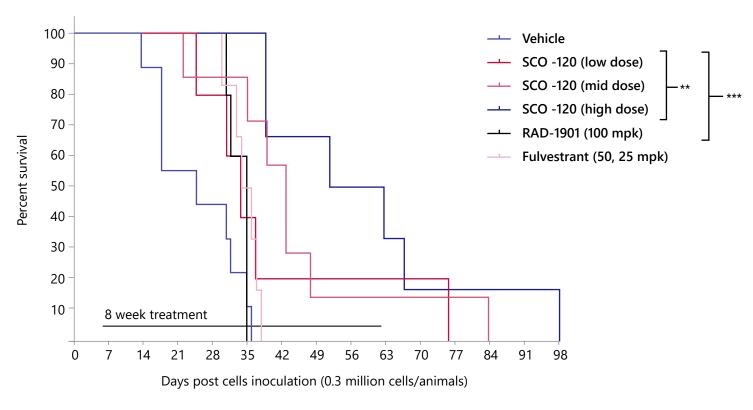


^{*}Fulvestrant group received 50 mg/kg as loading dose thrice- weekly for first week, followed by 25 mg/kg twice weekly for remaining 3 weeks | *p < 0.01 | ****p < 0.0001 as compared to vehicle treated group n.s.-non significant

SCO-120 advantage in brain metastases



Prolonged survival in preclinical brain-metastasis model expressing wild type $\text{ER}\alpha$

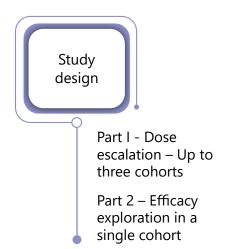


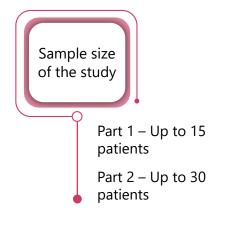
- Effectively crosses blood-brain barrier with higher accumulation in brain and tumor compared to plasma
- SCO-120 treated mice showed significant increased survival compared to RAD-1901 and fulvestrant
- Operation Potential to be an active treatment for HR+/HER2- breast cancer patients with brain metastases

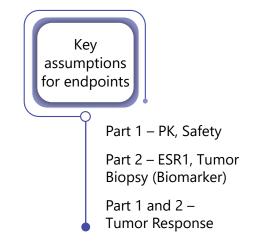
SCO-120 enters patient trials in 2022

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Clinical development plan and upcoming milestones









Concluding Remarks

Company highlights



Successful Track Record of Development and Commercialization along with a Robust Pipeline



USFDA approved drugs (Xelpros[™], Elepsia[™])



Indications targeted through 4 NCEs under clinical development



Preclinical programs in R&D pipeline covering 3 therapeutic areas

Targeting High Value Opportunities



USD 20Bn+

Combined peak sales potential for NCEs currently under clinical development



6 Licensing partners¹

Through an Innovationfocused R&D Platform with an Efficient Cost Structure



350+

Scientists across 4 research centers. Growing presence in the US (Princeton, NJ)



250+

Years of experience of management



Ongoing collaborations with universities / companies

^{1.} Licensing partners include Bioprojet, CMS, Sun Pharmaceutical Industries Ltd. (Sun Pharma), Tripoint Therapeutics, Biomodifying, and Visiox.

Highly experienced management team with global experience





















Nitin Damle Chief Innovation Officer Leads the development of Biologics Past experience:









Siu-Long Yao Head, Clinical Development & Operations Oversees design & execution of clinical research globally

Past experience:











Chetan Rajpara Chief Financial Officer

Responsible for finance, accounts, taxation and legal & secretarial functions

Past experience:









Nitin Dharmadhikari

Head, Operational Excellence & COEs

Responsible for New Initiatives, management of COEs and QA

Past experience:



Dr.Reddy's RANBAXY







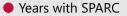
Head, Drug Discovery

Oversees Medicinal Chemistry, In-Vitro Biology, Bio-informatics & Process Development

Past experience:









Highly experienced management team with global experience











Head, Program Management

Oversees all aspects of the development / implementation of projects and programs Past experience:









Head, Drug Delivery Systems

Responsible for drug formulation and analytical development

Past experience:





Head, Translational Development

Responsible for Preclinical Pharmacology, Drug Metabolism & PK and Bioanalysis, and Regulatory Toxicology

Past experience:















Rajesh Ranganathan Head, Partnerships and Portfolio Strategy

Oversees external partnerships and portfolio management

Past experience:









Shanta Gupta

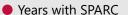
Chief Human Resource Officer

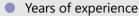
Responsible for the organization's human capital management

Past experience:

Reliance Infrastructure

STANTON CHASE





Scientific advisory board consisting of globally recognized experts





Phil Needleman, PhD Washington University in St. Louis





Rakesh Jain, PhD Massachusetts General HospitalHospital





Robert Spiegel MD, FACP¹ Weill Cornell Medical College, PTC Therapeutics









Mark Simon, MBA² Torreya Partners, Citigroup, Robertson Stephens, Kidder Peabody







Alan Ashworth, PhD, FRS **UCSF** ICR London





Jorge Cortes, MD Medical College of Georgia MD Anderson







Adrian Ivinson, PhD DRI UK, Nature, Harvard Medical School









Charbel Moussa, MBBS, PhD Georgetown University



Established and supported by marquee industry leader

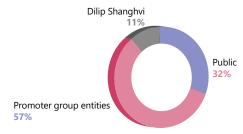




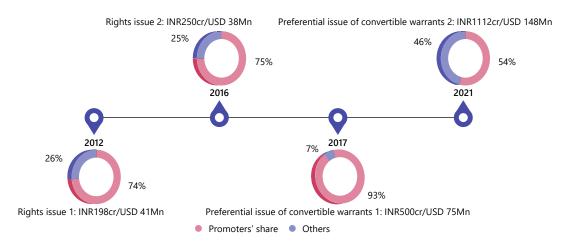
Dilip Shanghvi Chairman

- Founded Sun Pharma in 1983.
 (Current market cap of USD 24Bn+*)
- Has 35+ years of industry experience
- Awards and recognitions:
 Padma Shri (Fourth highest civilian award by Govt. of India) in 2016, Forbes Entrepreneur of the year 2014, Economic Times Business Leader of the Year (2014), CNN IBN's Indian of the Year (Business) (2011) and Ernst and Young's World Entrepreneur of the Year (2011).

Shareholding (as on 30th Sep. 2021)



Providing continuous support and investments



- Completed preferential issue for INR 1112 Cr. (USD 148Mn) in July 2021
- Well-capitalized for prosecuting the current clinical portfolio



Thank You

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