

SPARC/Sec/SE/2021-22/079

December 28, 2021

National Stock Exchange of India Ltd., Exchange Plaza, 5th Floor, Plot No. C/1, G Block, Bandra Kurla Complex, Bandra (East), Mumbai – 400 051. **BSE Limited,** Market Operations Dept. P. J. Towers, Dalal Street, Mumbai - 400 001.

Ref: Scrip Code: NSE: SPARC; BSE: 532872

Dear Sir/Madam,

Sub: Investor Presentation

Pursuant to Regulation 30 of the SEBI (Listing Obligations and Disclosure Requirements) Regulations, 2015, we enclose herewith the presentation, which the Company will be using at the meeting(s) to be conducted with the investors from today i.e. December 28, 2021 onwards.

The said presentation will also be uploaded on the Company's website, after sending this letter to you.

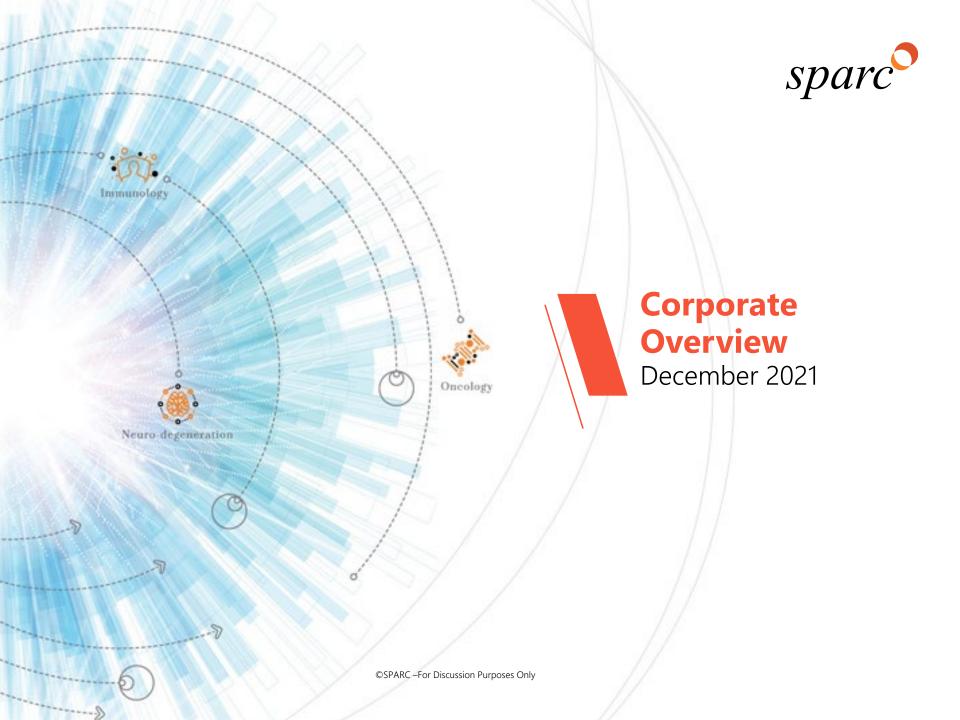
This is for your information and dissemination.

Yours faithfully,

For Sun Pharma Advanced Research Company Ltd.

Dinesh Lahoti Company Secretary and Compliance Officer ICSI Membership No. A22471

Encl: As above



Disclaimer



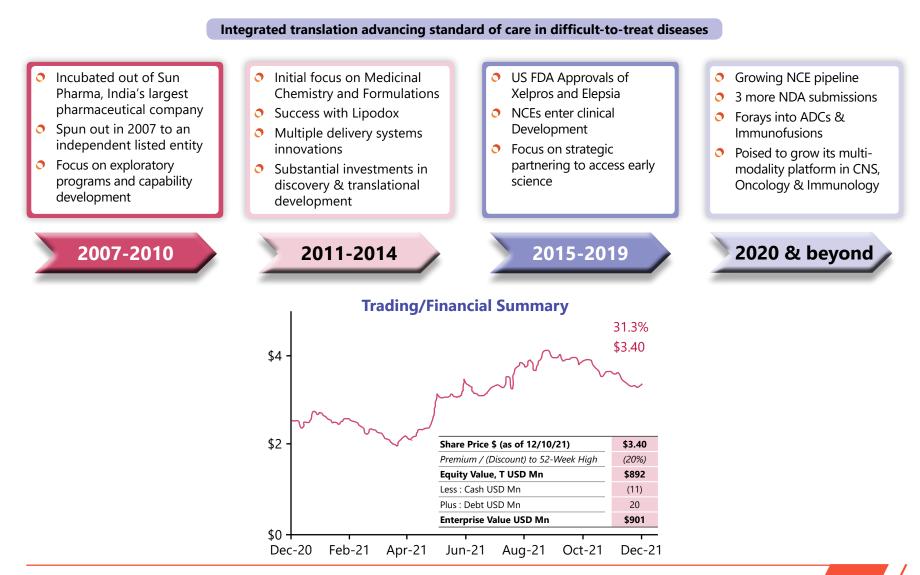
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SPARC timeline

Built a robust R&D engine over 15 years





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



Histol Myers Squibb





ADC = Antibody Drug Conjugate | TA = Therapeutic Area | USFDA = United States Food and Drug Administration | NDA = New Drug Approval

b NOVARTIS

Schering-Plough Wyeth

Pipeline overview & key milestones



Asset / Program	МоА	Indication	Discovery	Preclinical	Phase 1	Phase 2	Phase 3/ Registration Study	Upcoming Catalyst	Partner
		Parkinson's Disease						PoC data from PROSEEK study in 2023	
Vodobatinib (SCC-138)	c-ABL Inhibitor	Lewy Body Dementia ¹						PoC data in 2023	
		Alzheimer's Disease							
Vodobatinib (SCO-088)	BCR-ABL Inhibitor	Refractory CML						Pivotal data in 2024	
SCO-120	SERD	Metastatic Breast Cancer						Phase 1 data in 2023	
Vibozilimod (SCD-044)	Selective S1PR1 agonist	Psoriasis						Phase 2 data in 2023	SUN PHARMA
		Atopic Dermatitis						Phase 2 data in 2023	SUN PHARMA
			Alopecia Areata						
Undisclosed	TAA-1	Multiple Tumors						IND Filing Targeted 2023	
Preclinical Assets	10+ preclinio	cal assets under	development	to ensure a ro	bust pipeline	e for future g	Irowth		
			Neurol	ogy 🛑 C	Oncology	lmmun	ology		

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



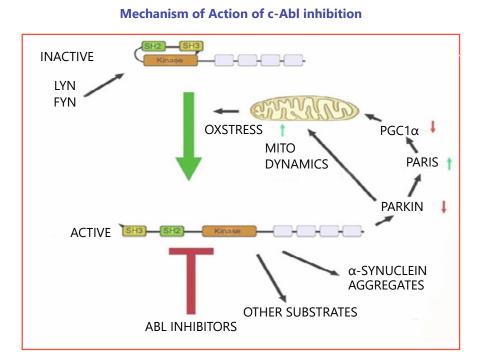
Vodobatinib for Neurodegenerative Diseases (SCC-138)

A potential first-in-class disease modifying therapy

Vodobatinib for neurodegenerative diseases

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Optimal agent to test the c-Abl hypothesis



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

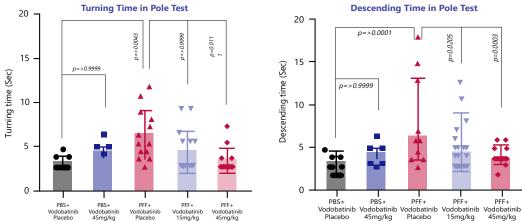
Family	Kinases	IC ₅₀ (nm)	
Abl	Abl (Abl-1)	0.9	
ADI	Arg (Abl-2)	0.8	
	Src	90.0	
	Fyn	18.0	
	Hck	54.0	
SFK	Lck	17.0	
	Lyn	18.0	
	Yes	28.0	
	PTK5	3.0	

Behavioral assessments in the PFFinduced mouse model

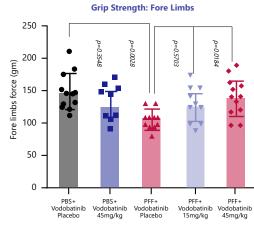


- In the MPTP¹ mouse model, Vodobatinib prevents neuronal degeneration in substantia nigra
- In the PFF² induced mouse model, vodobatinib shows target engagement, reduction in Serine 129 phosphorylation of α-Synuclein, preservation of dopaminergic neurons and clinical improvement in motor and cognitive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

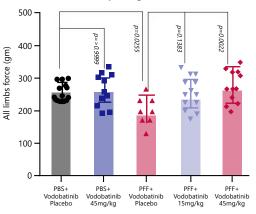
Vodobatinib at 45 mg/kg improves PFF-induced movement disorderrelated deficits in Turning Time and Descending Time in the Pole test



Vodobatinib treatment improves PFF-induced deficits in Grip Strength

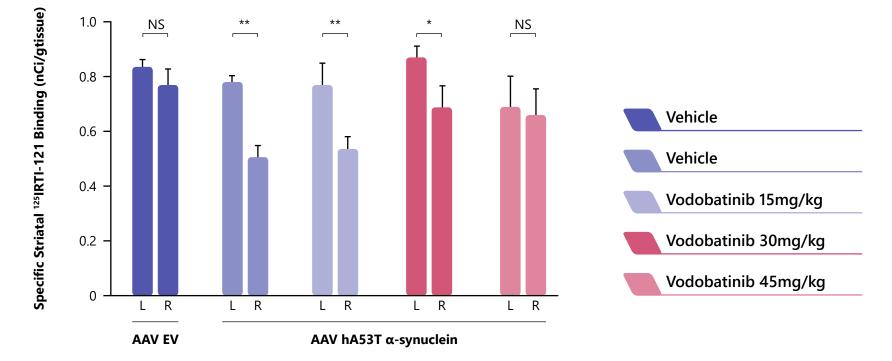


Grip Strength: All Limbs



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

Study conducted by Atuka Inc. | Unpublished data; not to be replicated or shared.

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Vodobatinib met the brain exposure targets in early clinical studies

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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

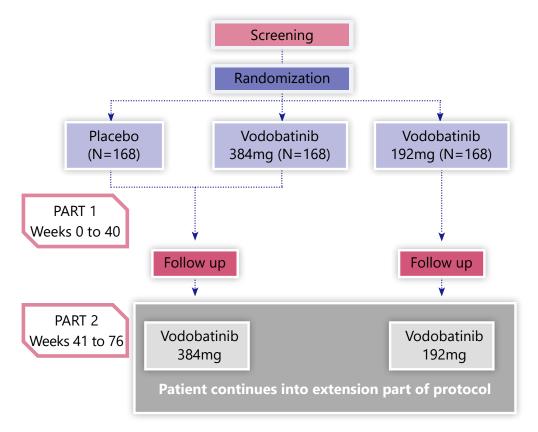
- Phase 1 completed in healthy subjects, PD subjects up to 384mg
 - Overall well tolerated
 - PK suggests adequate brain penetration over 24 hours

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

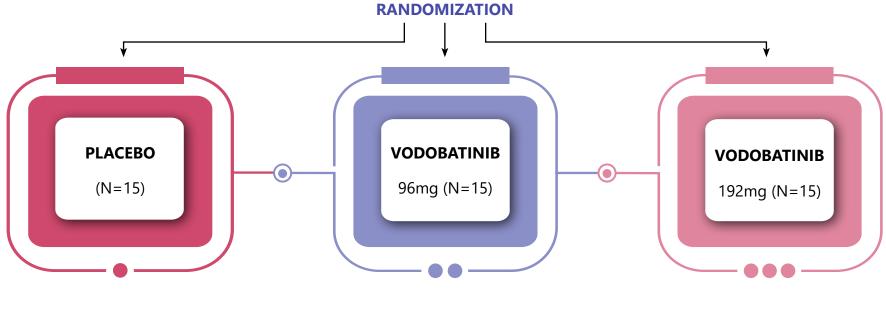
Data cut-off date: 26th Nov 2021 | A Phase 2 Study In Early Parkinson's Disease Patients Evaluating The Safety And Efficacy Of Abl Tyrosine Kinase Inhibition Using K0706 K0706 = Vodobatinib PD (SCC-138) | NCT03655236

Opportunities beyond Parkinson's Disease



Dementia with Lewy Bodies offers an immediate next opportunity

- DLB 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



*https://ghr.nlm.nih.gov/condition/dementia-with-lewy-bodies



Vibozilimod (SCD-044) -A Selective S1PR1 Agonist

A safer alternative to JAK inhibitors

Vibozilimod (SCD-044)

disorders, particularly in dermatology



An opportunity to improve oral standard of care in dermatology

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



blockade

S1PR1 agonists	EC ₅₀			
	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

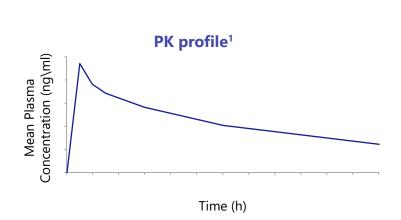
- Six dose levels in males and one dose level in females
- ~55% lymphocyte count decrease following 1 mg dose

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



%



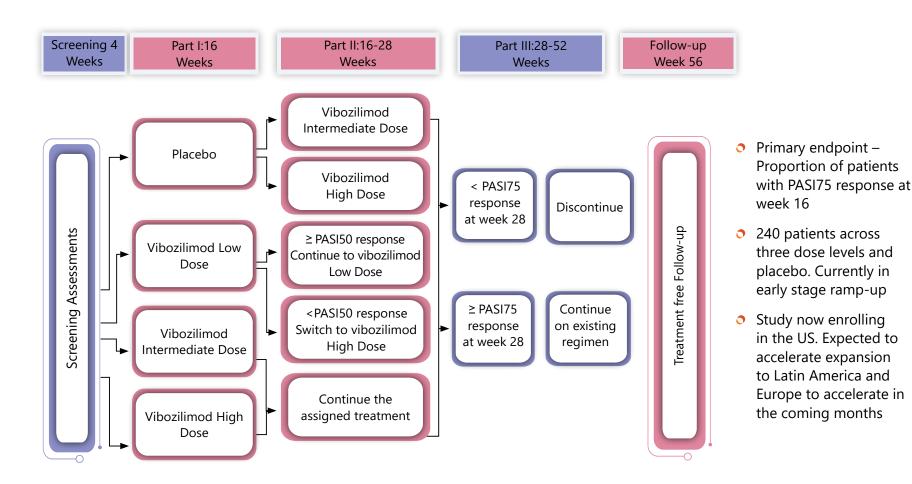
Lymphocyte count reduction¹



Vibozilimod (SCD-044) for psoriasis



Clinical proof-of-concept by 2023

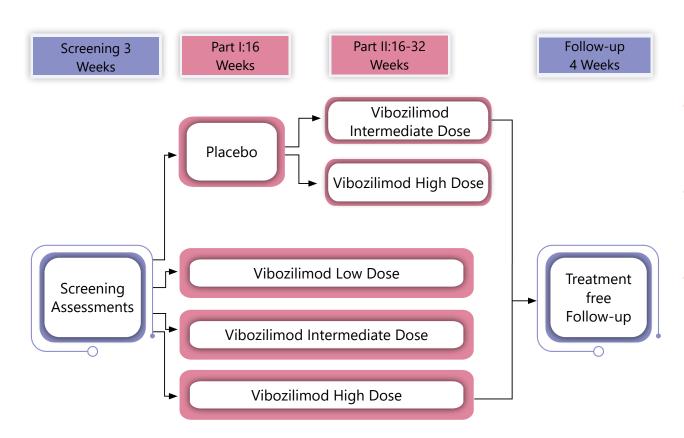


PASI = Psoriasis Area Severity Index | NCT04566666 (SCD-044-19-14, Version 1, September 23, 2020) | Vibozilimod (SCD-044) licensed to Sun Pharmaceutical Industries Limited

Vibozilimod (SCD-044) for atopic dermatitis



Clinical proof-of-concept by 2023



- Primary endpoint –
 Proportion of patients with EASI-75 response at week 16
- 240 patients across three dose levels and placebo. Currently in early stage ramp-up
- Study now enrolling in the US. Expected to accelerate expansion to Latin America and Europe to accelerate in the coming months

EASI = Eczema Area Severity Index | POC = Proof of Concept | NCT04684485 (SCD-044-19-16, Version 1, November 9, 2020) | Vibozilimod (SCD-044) licensed to Sun Pharmaceutical Industries Limited

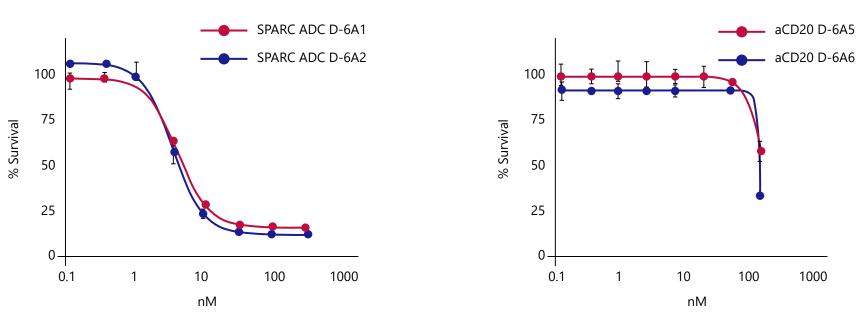


Anti TAA-1 Asset

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

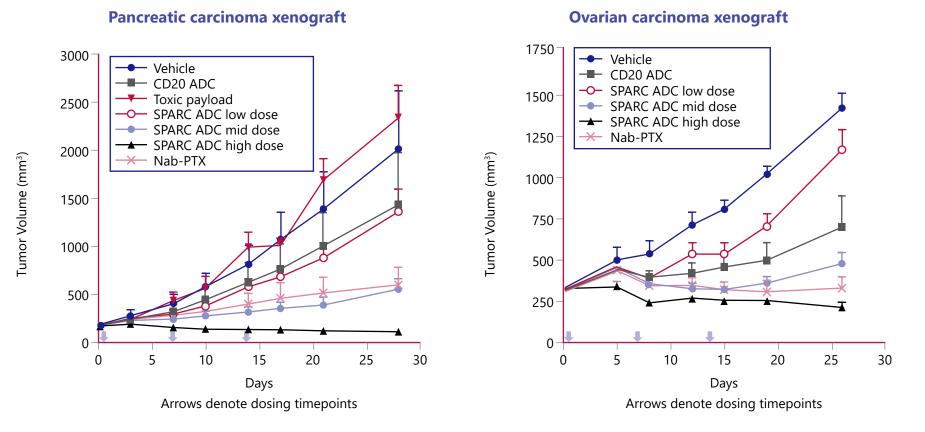
- ADC against a novel tumor associated antigen as a target
- Evidence of potent cytotoxicity of SPARC ADC against TAA-1 over-expressing pancreatic carcinoma cell line
- 100-fold greater potency over a nonbinding ADC of the same payload targeted to CD20

ADC = Antibody Drug Conjugate | TAA-1 = Tumor Associated Antigen-1 | CD20 = Cluster of differentiation 20

Antitumor efficacy of SPARC ADC

Efficacy established in multiple xenograft models





- o 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

ADC = Antibody Drug Conjugate | Nab-PTX = Nanoparticle albumin-bound Paclitaxel

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)



Promising Last Line Therapy

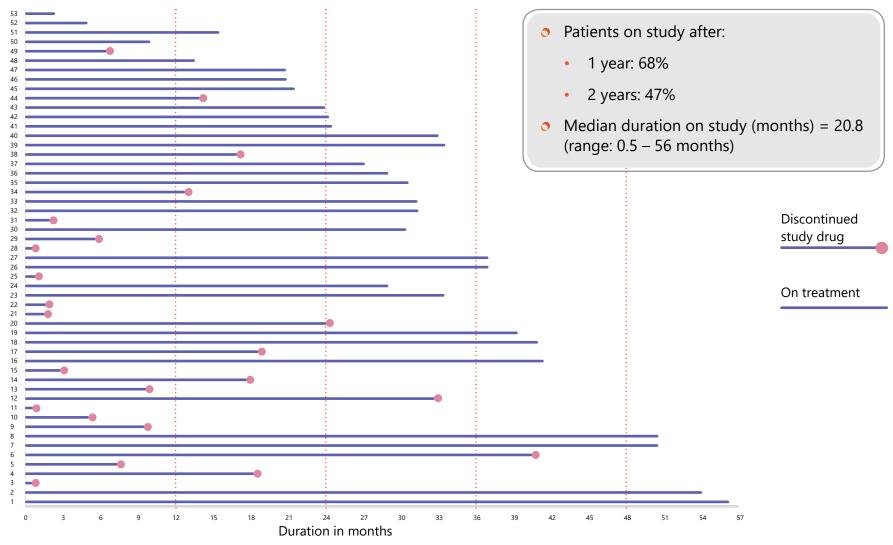
CML Overview	Market Opportunity	Product Overview	Clinical Summary
• CML is caused by a translocation of the abl gene that results in formation of Philadelphia Chromosome	• Branded 2nd and 3rd generation TKIs retain high commercial value due to refractory nature of CML, despite genericization of 1st generation TKI	• Targeting patients who are refractory and/or intolerant to other TKIs	 Phase 1 completed in CML subjects
 Prior to the discovery of BCR-ABL inhibitors, CML was a fatal disease with an 8-year survival rate of ~6% 	 Large market opportunity – US drug sales of the CML TKIs over \$3Bn² 	 Well tolerated with significant coverage of the mutational field 	• Favorable safety and tolerability
• Tyrosine kinase inhibitors have changed the prognosis of CML, but patients eventually can become resistant to drugs	 O 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 	 Has shown promising activity in clinical trials 	 Registration study underway. Planned US NDA filing in 2024
 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 	 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 	 Orphan Drug Designation and Accelerated Approval pathway agreed with USFDA 	

TKI = Tyrosine Kinase Inhibitor | 1. SEER database Cancer Stat Fact | 2. IQVIA 2021

Vodobatinib for CML (SCO-088)

Durable long-term responses seen across cohorts





Data cutoff 29th November 2021 | Unpublished data, not to be replicated | Number on Y-axis represents individual patients

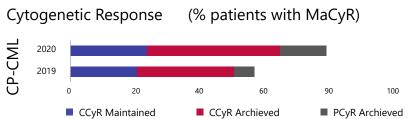
Vodobatinib for CML (SCO-088)



Clinical Development Plan

Pivotal (Part C) study ongoing

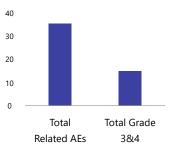
	Clinical Development Plan					
Part A	Single Ascending Dose study (SAD) in volunteers	\odot	 Orphan Drug Designation approved by USFDA and EMA Market exclusivity in addition to IP coverage User fee waiver 			
Part B	Multiple Ascending Dose study (MAD) in patients	\odot	 EOP1 discussion completed; agreement with USFDA reached on accelerated approval pathway based on 			
Part C	Pivotal efficacy study in refractory and/or intolerant patients to 3 prior TKIs		Part C (pivotal study)			
	Efficacy	Safety and Tolerability				



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

 Generally well tolerated with slight excess of GI and hematological AEs

All Treatment Emergent AEs (Cases)



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 Overview







- Breast cancer is the second most common cancer diagnosed in women in the United States¹
- Annual incidence of ~2 million patients across the world¹

0

- ~70% of the breast cancer is HR+/HER2-1
- Hormonal therapy is SoC for ~70% of HR+/HER2metastatic breast cancer patients1. 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

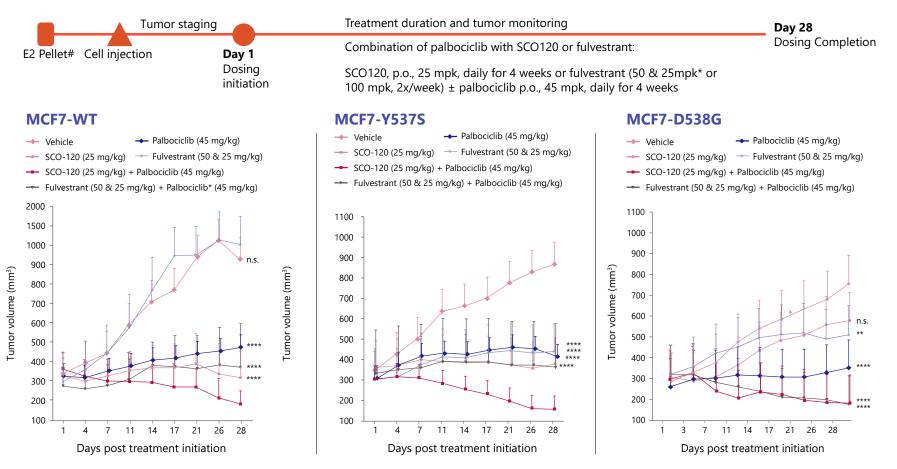
1. CancerMPact® Treatment Architecture U.S., Breast Cancer | HR = Hormone Receptor | HER2 = Human Epidermal Growth Factor Receptor 2 | ER α = Estrogen Receptor α | SOC = Standard of Care | IM = Intramuscular SERD = Selective Estrogen Receptor Degrader | AE = Adverse Event | SERM=Selective Estrogen Receptor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

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In vivo efficacy of SCO-120 in combination with palbociclib



Promising activity against resistant mutants alone and in combination with palbociclib



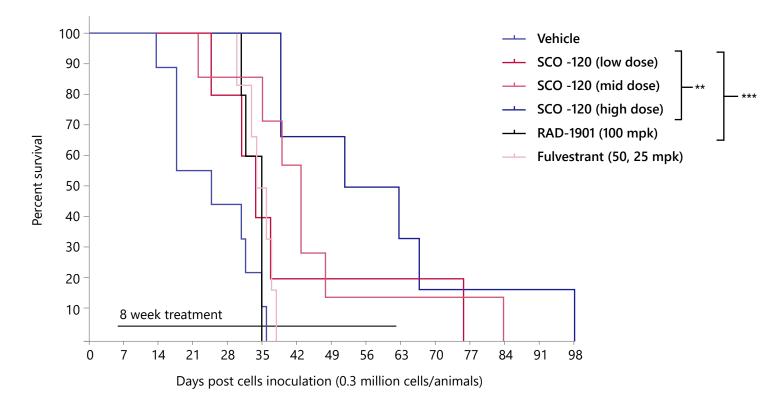
*Fulvestrant group received 50 mg/kg as loading dose thrice- weekly for first week, followed by 25 mg/kg twice weekly for remaining 3 weeks | *p < 0.01 | ****p < 0.0001 as compared to vehicle treated group n.s.-non significant

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SCO-120 advantage in brain metastases



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



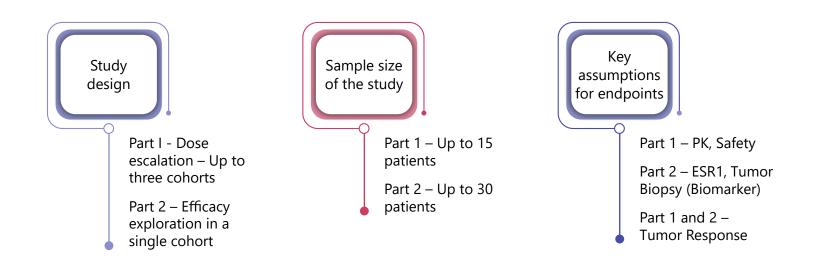
- Effectively crosses blood-brain barrier with higher accumulation in brain and tumor compared to plasma
- SCO-120 treated mice showed significant increased survival compared to RAD-1901 and fulvestrant
- Potential to be an active treatment for HR+/HER2- breast cancer patients with brain metastases

PK = Pharmacokinetic | ESR1 = Estrogen receptor 1

30

SCO-120 enters patient trials in 2022

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



Combined peak sales potential for NCEs currently under clinical development



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



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



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





Anil Raghavan Chief Executive Officer Responsible for strategic prioritization and portfolio decisions

Past experience:





Chetan Rajpara Chief Financial Officer

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

Past experience:





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



endo



Nitin Dharmadhikari Head, Operational Excellence & COEs Responsible for New Initiatives, management of COEs and QA Past experience:



Years with SPARC • Years

Years of experience



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

Past experience:





Trinadha Rao Chitturi Head, Drug Discovery

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

Past experience:



Highly experienced management team with global experience





Vikram Ramanathan 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

20

Past experience:



NOVARTIS



Shravanti Bhowmik Head, Program Management

Oversees all aspects of the development / implementation of projects and programs

Past experience:





Shanta Gupta Chief Human Resource Officer Responsible for the organization's human capital management Past experience:

Reli STANTON CHASE Infrastructure

Years with SPARC Years of experience



Yashoraj Zala Head, Drug Delivery Systems

Responsible for drug formulation and analytical development

Past experience:





Scientific advisory board consisting of globally recognized experts





Phil Needleman, PhD Washington University in St. Louis

Washington" University in St. Louis School of Medicine



Rakesh Jain, PhD Massachusetts General HospitalHospital



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

Schering-Plough

P



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



Alan Ashworth, PhD, FRS UCSF ICR London





MGH

Jorge Cortes, MD Medical College of Georgia MD Anderson



Weill Cornell

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





Charbel Moussa, MBBS, PhD Georgetown University



Established and supported by marquee industry leader

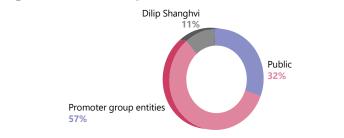




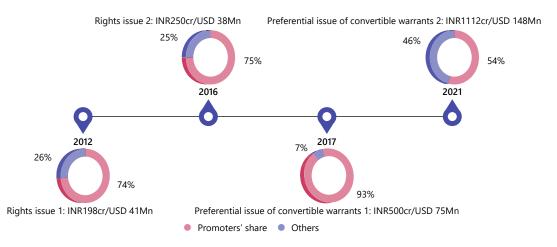
Dilip Shanghvi Chairman

- 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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