



Pioneering Innovative Treatments for Fibrotic Diseases

Corporate Overview | Non-Confidential | January 2022

Forward Looking Statements



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Investment Highlights

Clinical Stage Biotech Company Focused on Inflammatory-Fibrotic Diseases



Focus

- Discovery and development of innovative therapeutics for rare inflammatory and fibrotic diseases with high unmet need

CM-101: A Pipeline in A Product

- CM-101, a first-in-class CCL24 neutralizing monoclonal antibody targeting:
 - Primary Sclerosing Cholangitis (PSC)
 - Systemic Sclerosis (SSc)
 - Liver fibrosis (MoA*)

Near-Term Catalysts

- Conducting 2 Phase 2 clinical trials with 3rd Phase 2 trial expected to initiate in 1Q22
- First clinical readouts targeted for 2022 expected to drive multiple value inflections

Robust IP Portfolio

- Global patent exclusivity through 2041. Granted composition of matter patent with multiple additional patents pending

Top Tier Investors



**Cormorant Asset
Management**

THIEL

Solid Balance Sheet

- Cash & equivalents of \$64.3M as of September 30, 2021, enabling runway through mid-2023

*MoA – Mechanism of Action

Experienced Management



DALE PFOST, PhD

Incoming Chairman & Chief Executive Officer*

OPKO

**Advent
Life Sciences**

**SmithKline Beckman
CORPORATION**



ORCHID



ADI MOR, PhD

Co-founder & Chief Scientific Officer



TEL AVIV
UNIVERSITY תל אביב אוניברסיטת



DON MARVIN

Chief Financial Officer, EVP and COO



ORCHID



Abbott



BAYER



concentric
THE PHYTO-MICROBIOME COMPANY



Boehringer
Ingelheim



DAVID WEINER, MD

Incoming Interim Chief Medical Officer



ACADIA



EMD
SERONO



lumos
PHARMA



aTyr



PROTEOSTASIS



SHARON ELKOBI

VP Business Development



teva



MEDISON
Delivering Innovative Healthcare



Neurim
PHARMACEUTICALS



MICHAL SEGAL-SALTO, PhD

VP Research and Development



TEL AVIV
UNIVERSITY תל אביב אוניברסיטת



מכון ויצמן למדע
WEIZMANN INSTITUTE OF SCIENCE

Experienced Leadership



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Dale Pfof, PhD

*Incoming Chairman**
Chief Executive Officer



Neil Cohen, MA

Director



Nissim Darvish, MD, PhD

Director



Joel Maryles, CFA, MBA

Director



Adi Mor, PhD

Director
Co-founder & Chief Scientific Officer



Alan Moses, MD

Director



Claude Nicaise, MD

Director



Scientific Advisory Board (partial)

Prof. Marco Matucci-Cerinic, MD, PhD

Director of the Division of Rheumatology,
University of Florence, Italy

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Head of the Scleroderma Program at NIHR,
University of Leeds, UK

Scott L. Friedman, MD

Dean for Therapeutic Discovery and Chief,
Division of Liver Diseases, Mount Sinai, NY, USA

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Lily and Terry Horner Chair in Autoimmune Liver
Disease, University of Toronto, Toronto General
Hospital, Canada

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Director of the Scleroderma Program,
University of Michigan, Ann Arbor, Michigan, USA

Massimo Pinzani, MD, PhD, FRCP

Sheila Sherlock Chair of Hepatology, Director UCL
Institute for Liver and Digestive Health, RFH, London, UK

* Subject to shareholder approval

CM-101: Pipeline in a Product

Addressing Fibrotic Diseases with High Unmet Need

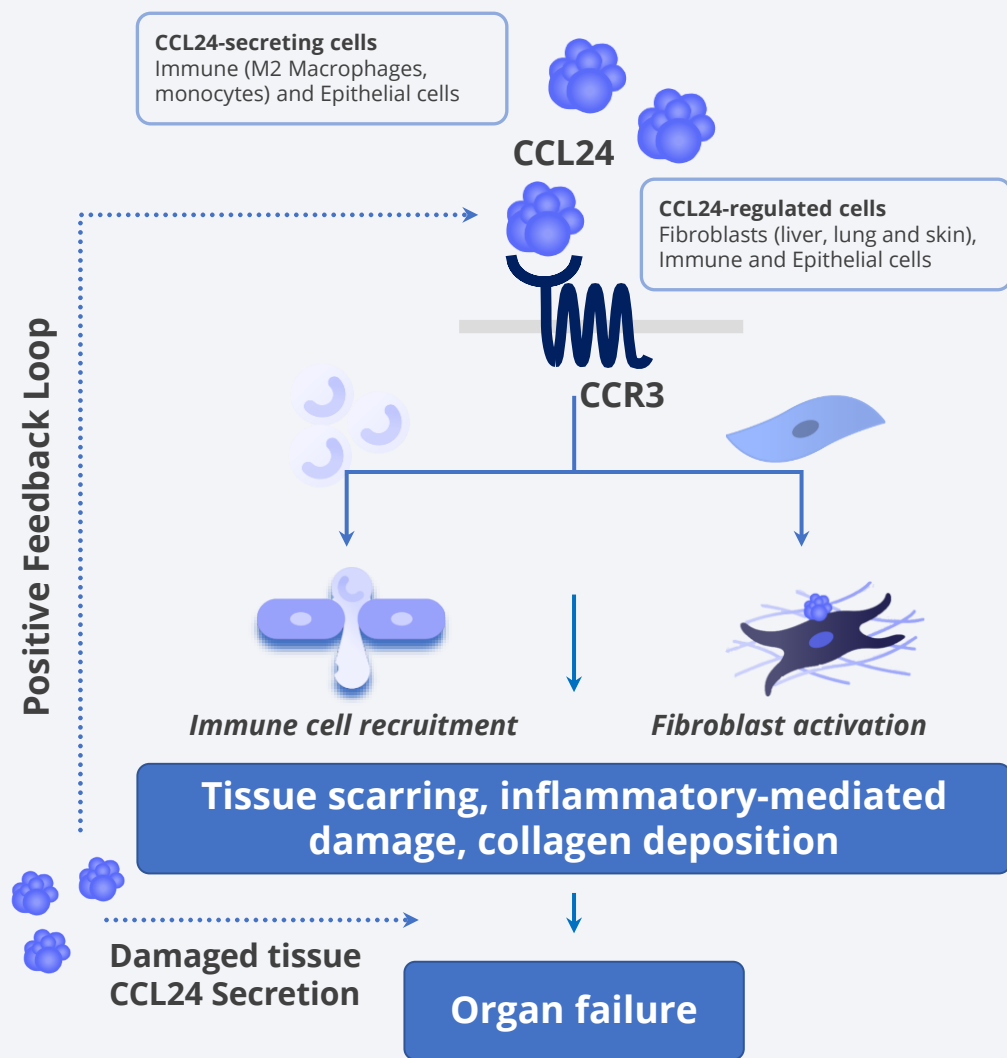


Program	Indication	Target	Discovery	Preclinical	Phase 1	Phase 2	Status
CM-101	Primary Sclerosing Cholangitis	CCL24	<i>Orphan designation granted from FDA and EMA</i>				Enrolling Patients Top Line Data 2H 2022
CM-101	Systemic Sclerosis	CCL24	<i>Orphan designation granted from FDA and EMA</i>				Initiation Expected In 1Q 2022
CM-101	Liver Fibrosis (MoA*)	CCL24	<i>Mechanism of Action & SC Route of Administration (in NASH patients)</i>				Enrolling Patients Top Line Data 1H 2022
Next-gen	Fibrosis	Screening					In Progress

*MoA – Mechanism of Action

CCL24 is a Novel Therapeutic Target for Fibrosis

Critical Mediator Promoting Inflammation and Fibrosis



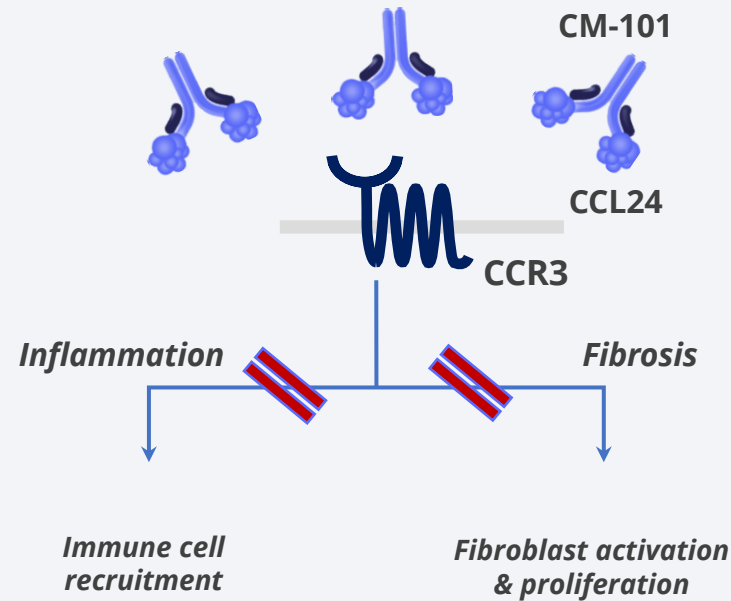
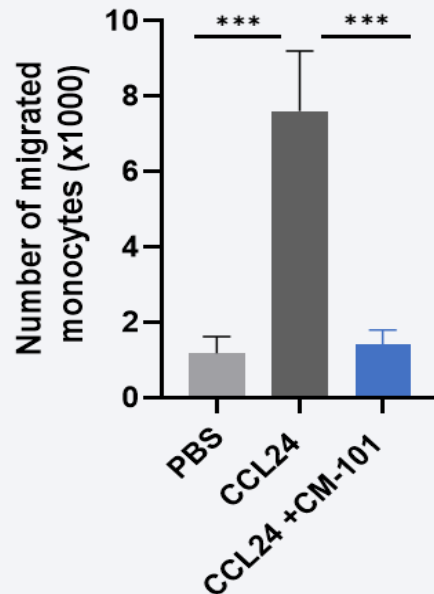
- ✓ **Dual role in promoting fibrosis**
 - directly activates fibroblasts
 - enhances local immune cell recruitment
- ✓ **Unique and differentiated activity**
 - *ex vivo* and *in vivo* data confirm unique role vs other CCLs
 - correlates with disease outcome and fibrotic biomarkers
- ✓ **Minor expression in healthy tissue**
 - significantly elevated in liver, skin, lung fibrotic tissue
 - wide therapeutic margin
- ✓ **Positive feedback loop potentiates tissue damage**
 - responsible for initiation and perpetuation of fibrosis

CM-101: A First-in-Class mAb Blocking CCL24

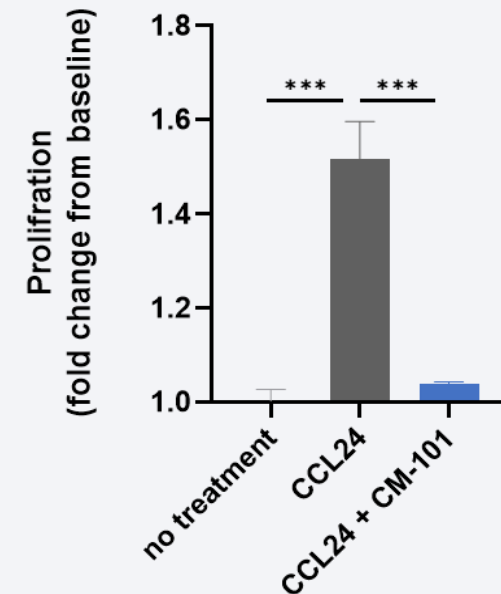
Dual Mechanism of Action Interfering with the Core Fibrotic Pathways



CM-101 Reduces *In-vivo* Monocyte Recruitment



CM-101 Inhibits Primary Hepatic Fibroblast Activation



CM-101 attenuates inflammation and fibrosis by inhibiting fibroblast activation and immune cell recruitment

Strong Preclinical Evidence of CM-101 Activity via CCL24 Inhibition

Significant Attenuation of Fibrosis & Inflammation Across a Wide Range of Models

CCL24 Target Validation

Ex Vivo (Patient Samples)

Primary sclerosing cholangitis

- Biomarker correlations
- Overexpression of CCL24 and CCR3

Systemic Sclerosis

- Fibrotic biomarkers correlation
- Disease deterioration correlation
- Overexpression of CCL24 and CCR3

Liver Fibrosis (NASH)

- Disease severity correlation
- Overexpression of CCL24 and CCR3

In Vivo (Knockout Animal Models)

Systemic Sclerosis

- CCL24 knock out vs. WT in Bleomycin induced skin fibrosis model (mice)

Liver metabolism and inflammation

- CCL24 knockout vs. WT in MCD-induced NASH (mice)

Proof of Concept\Animal Models

Primary sclerosing cholangitis

- ANIT induced cholestasis-chronic and acute (mice)
- Bile duct ligation (rat)
- MDR2 knock-out (mice)

Systemic sclerosis

- Bleomycin-induced skin fibrosis (mice)
- Bleomycin induced lung fibrosis (mice)

Liver fibrosis

- TAA-induced liver fibrosis (rat and mice)

Liver metabolism and inflammation

- STAM (mice)
- MCD diet induced NASH (mice)

Atherosclerosis

- ApoE knock out model (mice)

Mechanism of Action

CM-101 effects on fibroblast activation

- Dermal, hepatic and lung fibroblast activation
- Dermal and liver fibroblast transition to myofibroblast
- Hepatic fibroblast motility

CM-101 effects on immune cell migration and recruitment

- Dermal fibroblast migration
- Monocyte polarization
- Monocyte recruitment

Toxicology

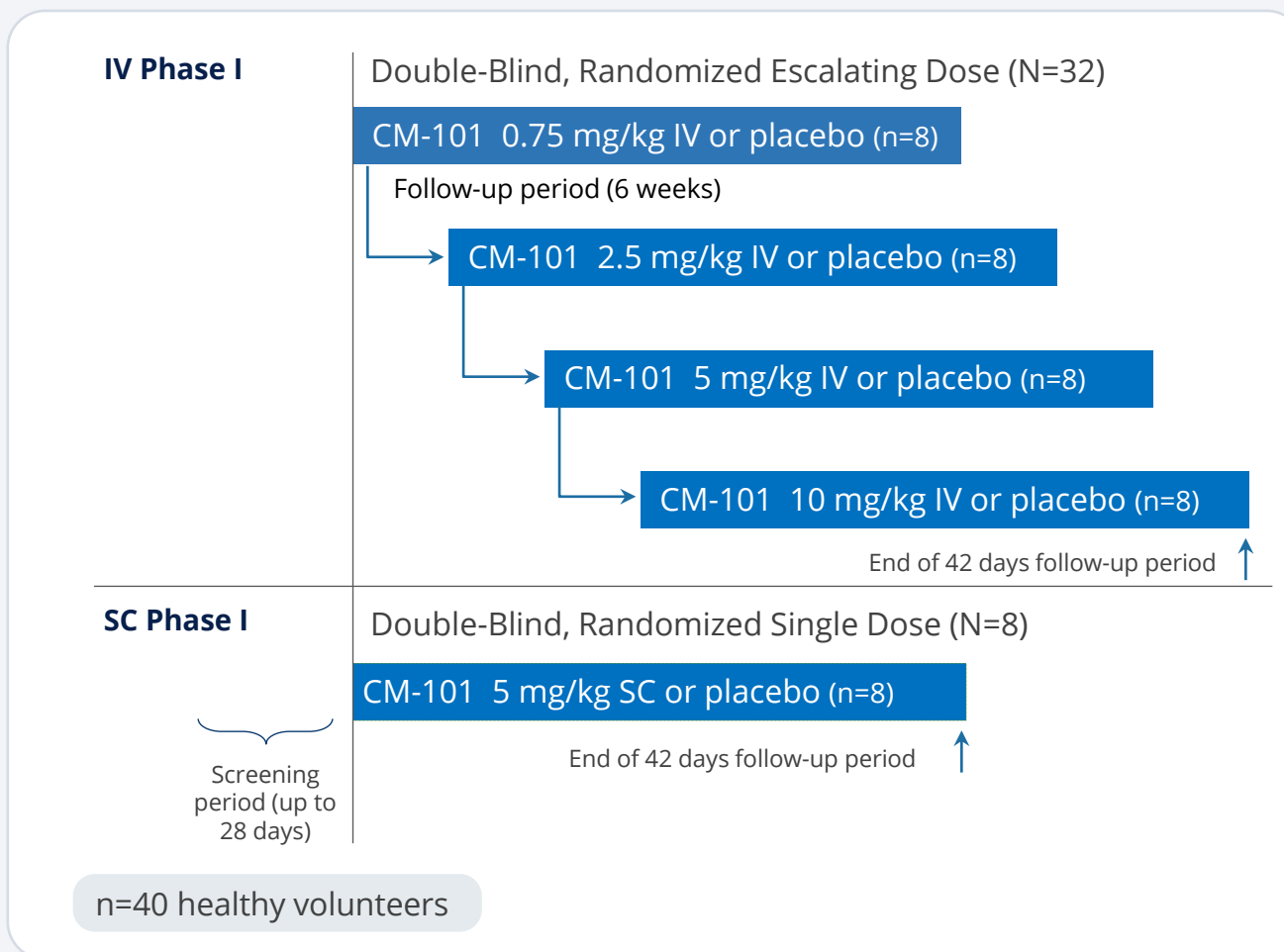
- **Short-term GLP** in rodents
- **Long-term GLP** in non-human primates
- **Ex vivo safety:** ADCC, CDC, cytokine secretion
- **Tissue cross reactivity**

Phase 1a Results



Single Ascending Dose Study Shows CM-101 to be Safe and Well Tolerated in Healthy Volunteers

- CM-101 was **safe and well tolerated** at all tested doses up to 10 mg/kg and for both formulations
- All drug related AEs were classified as **mild with no SAEs reported**
- Average half-life of 19-21 days (for IV and SC) supports long interval **administration once every 2-4 weeks**
- **Dose-dependent target engagement** measured by serum CCL24 levels
- **Comparable target engagement & PK profiles** for the SC and IV formulations



Phase 1b Results

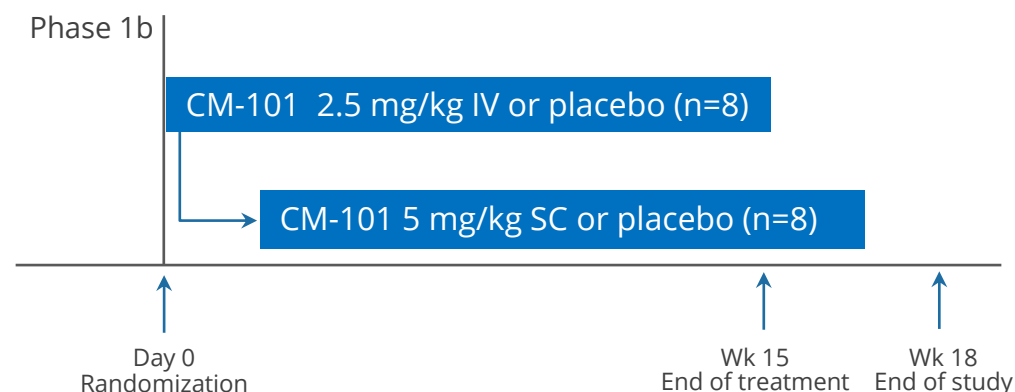
Multiple Administration Study Showed CM-101 to be Safe & Well Tolerated in NAFLD Patients



- Study population: **NAFLD patients** with normal liver function
- Multiple CM-101 administrations were **safe and well tolerated** using both **IV** and **SC** formulations
- Most frequently reported AEs were **mild** with **no drug-related SAEs reported**
- Favorable $t_{1/2}$ supports **long dosing interval** (Q2W - Q4W)

Study Design

Double-Blind, Randomized Escalating Dose (N=16)

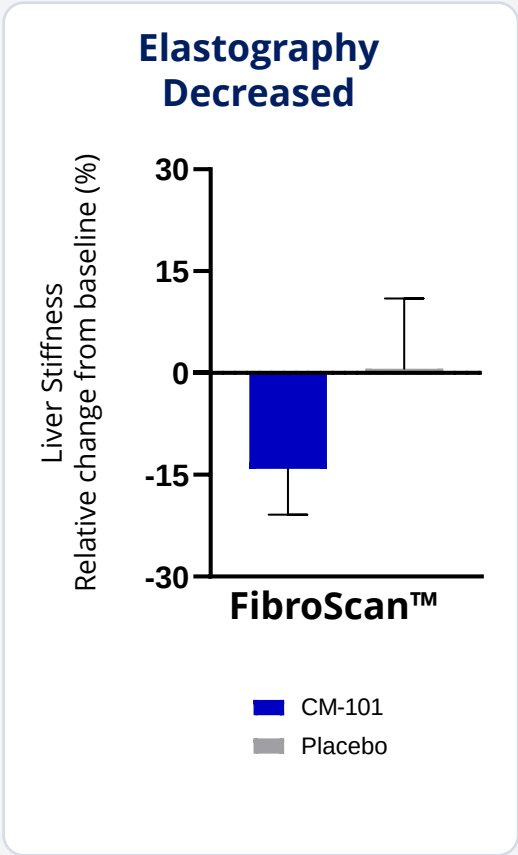
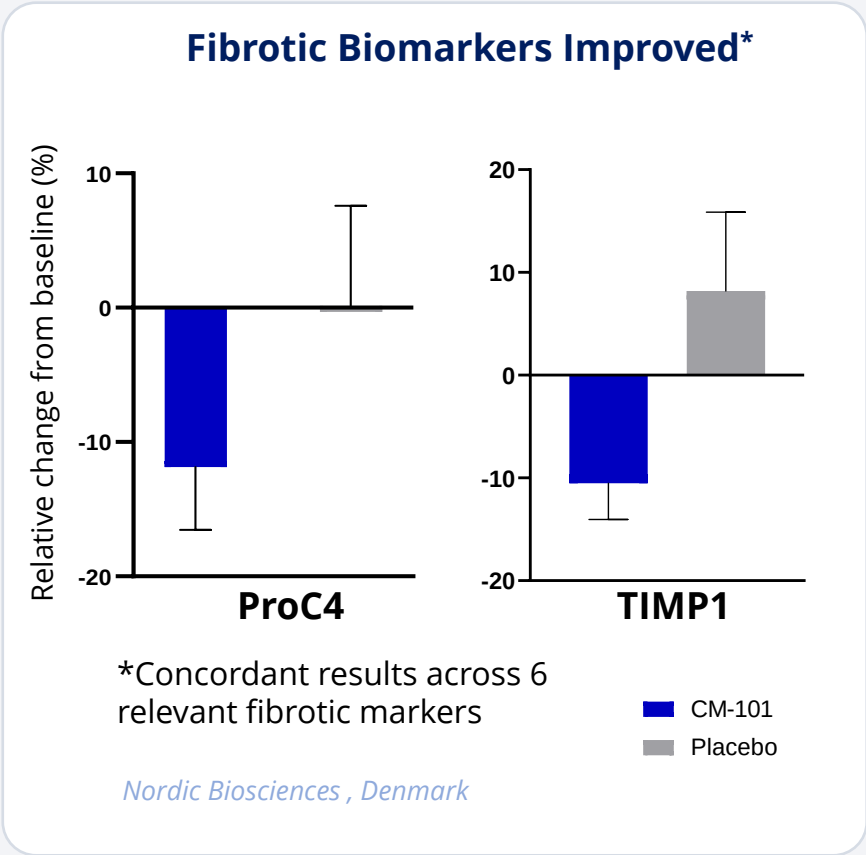
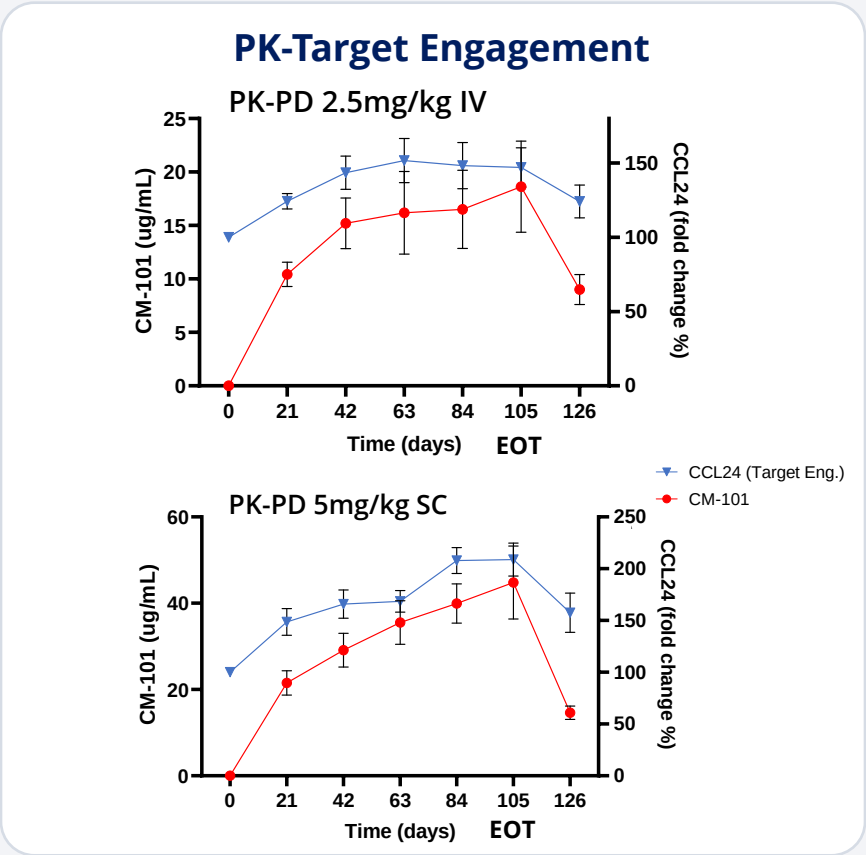


- Tested doses - 2.5 mg/kg IV infusion and 5 mg/kg SC injection
- 5 repeated administrations per patient; Q3W
- Primary endpoint - safety and tolerability

Phase 1b: Confirms CM-101 Anti-Fibrotic Mechanism in Humans



Demonstrated Biomarker Improvement with Dose-Dependent PK-Target Engagement



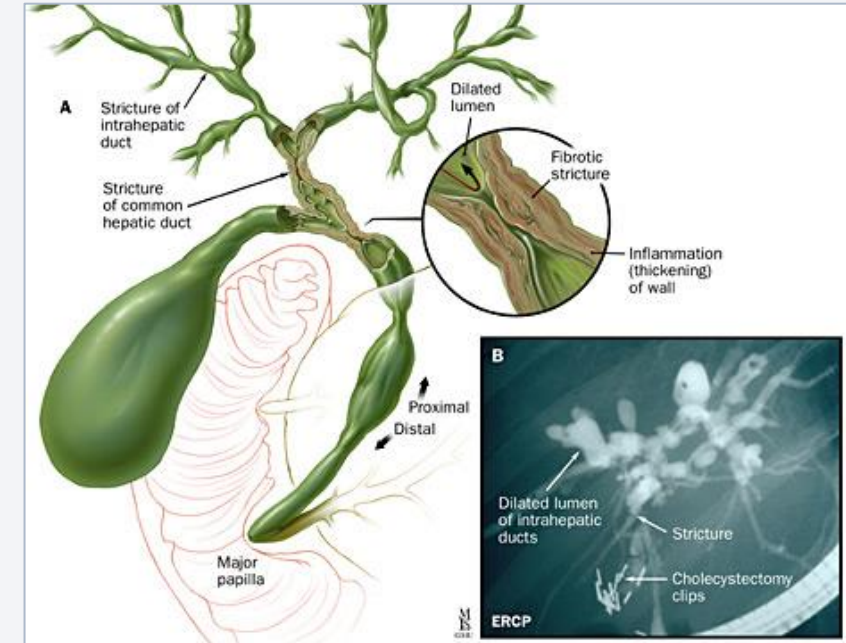
CM-101 for the Treatment of Primary Sclerosing Cholangitis



Primary Sclerosing Cholangitis (PSC)

Characterized by High Morbidity and Mortality

- **PSC is a rare, chronic liver disease** characterized by progressive course of cholestasis **with inflammation and fibrosis** of the intrahepatic and extrahepatic bile ducts **leading to end-stage liver disease and cirrhosis**
- Diagnosis is most often between the **ages of 30 - 40 years**; Predominantly affects men (2:1 male:female ratio)
- **No FDA approved drug**; Liver transplant is the only therapy with curative potential
- **Median survival of 10-12 years** with no intervention
- ~77K patients in 7 major markets



Source: Johns Hopkins Medicine

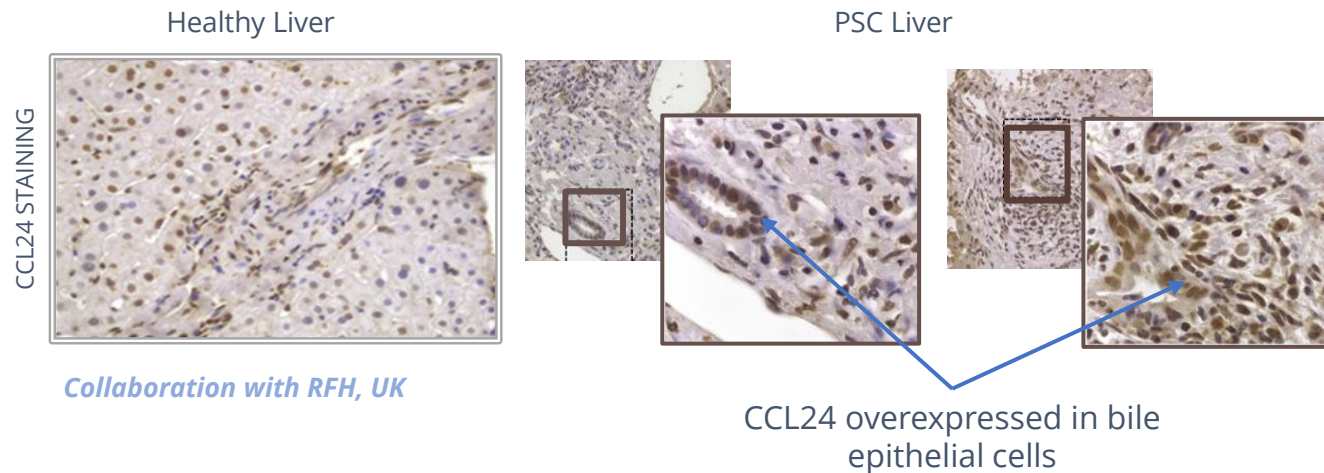
CCL24 Plays a Key Role in PSC

Clinical Relevance of CCL24 in PSC Pathophysiology is Supported by Patient Samples



CCL24 expression is significantly and selectively elevated in PSC livers

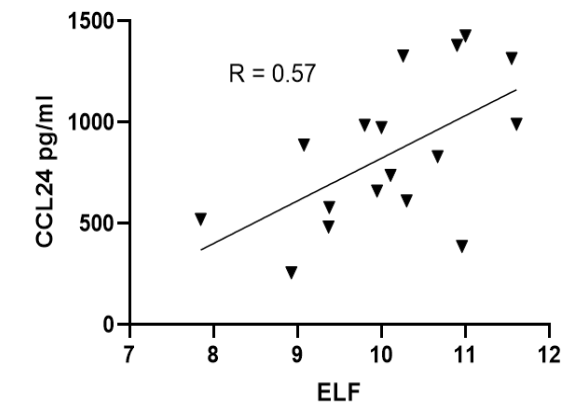
CCL24 level in healthy vs PSC patients' liver tissues



CCL24 correlates with ELF in PSC patients with elevated ALP

CCL24 serum level in PSC patients

ALP > 1.5 upper limit of normal



Enhanced Liver Fibrosis (ELF) test reflects fibrosis and predicts clinical outcomes in PSC¹

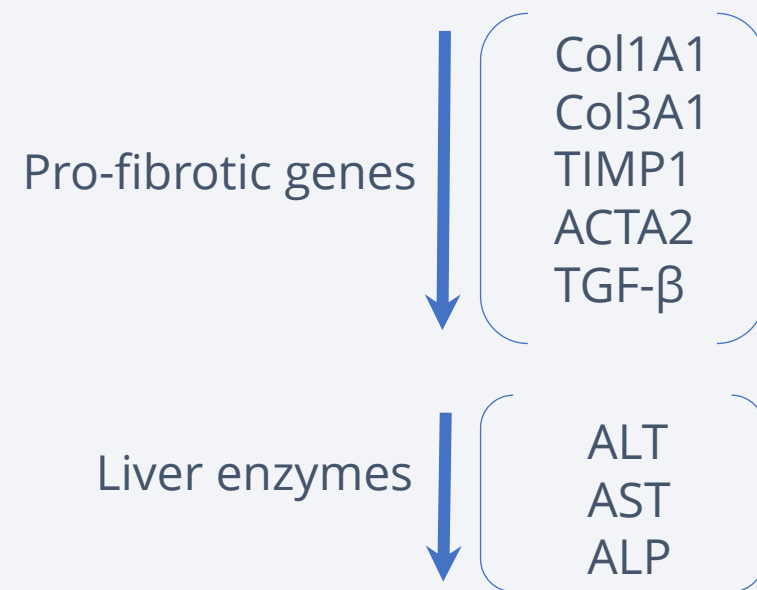
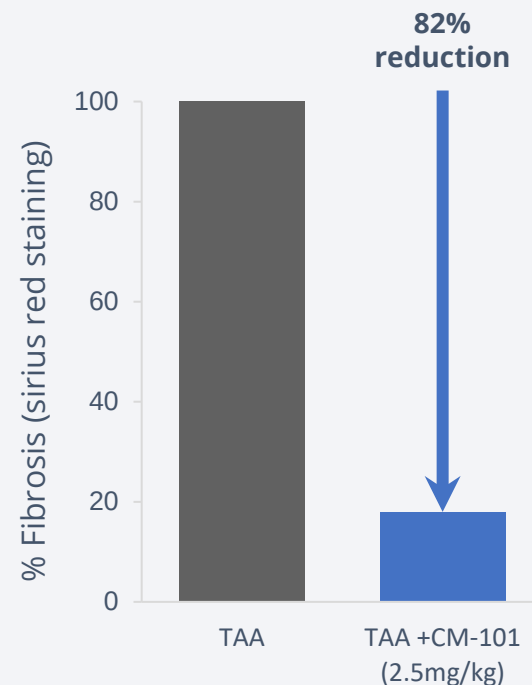
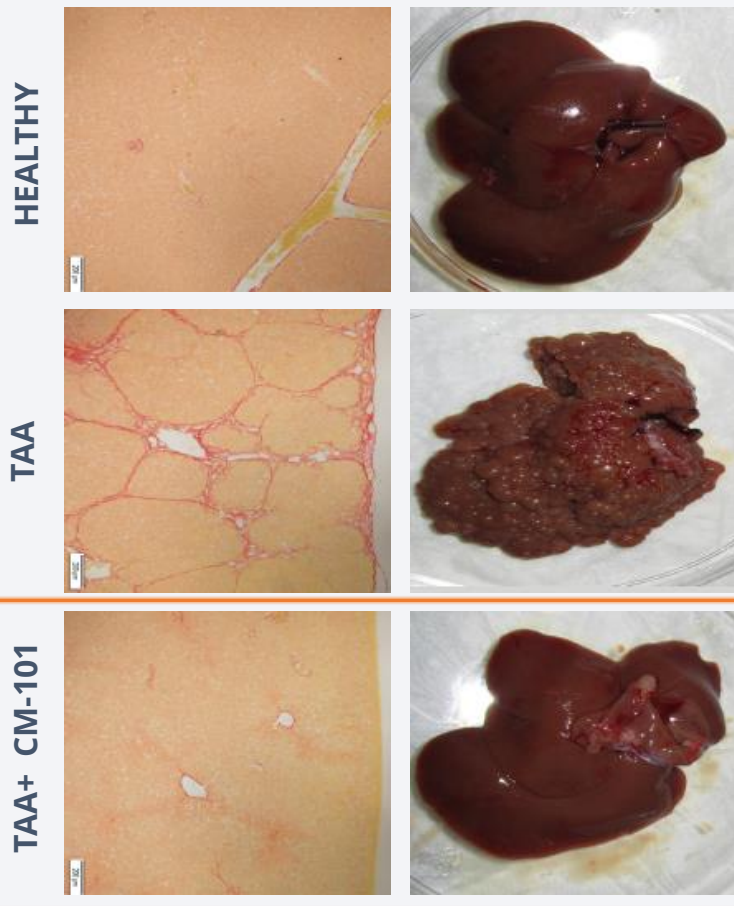
CM-101 MoA guided the selection of ELF and ALP as primary endpoints in Phase 2

CM-101 Reduces Liver Fibrosis by 82%

Reduced Liver Collagen in TAA Liver Fibrosis Rat Model Using Therapeutic Design

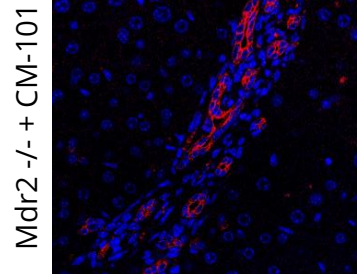
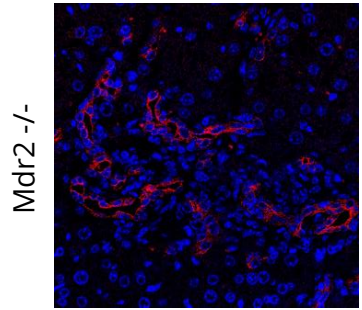


SIRIUS RED (COLLAGEN)



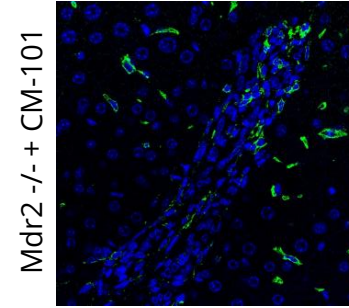
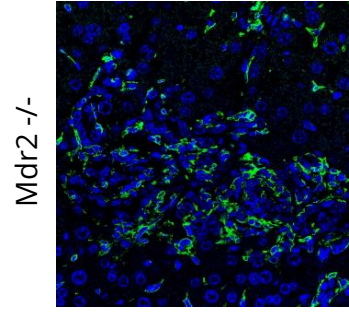
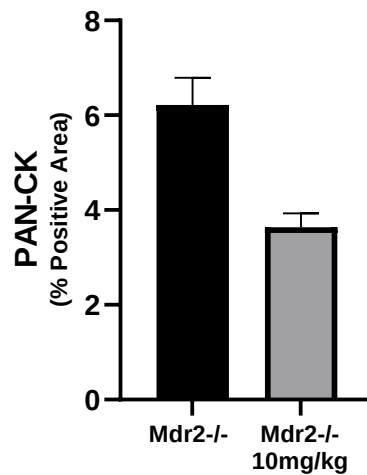
CM-101 Ameliorates Liver Injury and Fibrosis in PSC

Cholestasis, Inflammation and Fibrosis are Reduced in the MDR2 Knockout Model in Mice



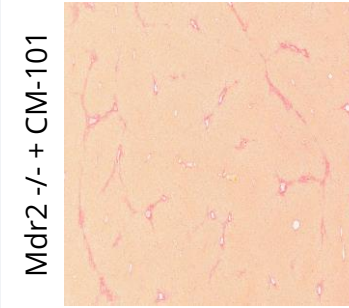
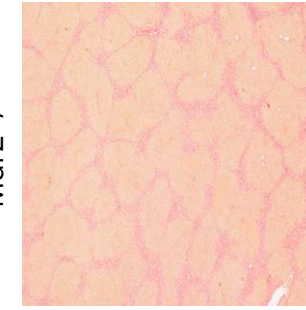
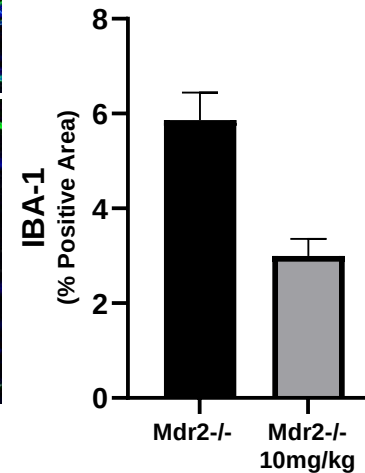
Cholangiocytes
(panck)

**Decreased Bile
Epithelial Cell
Proliferation**



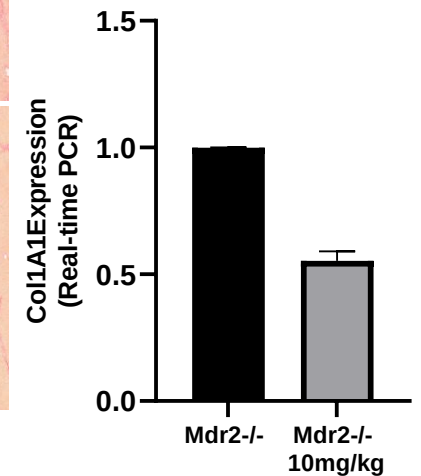
Macrophages (IBA1)

**Reduced
Macrophage
Number**



Sirius Red

**Attenuated Liver
Collagen Deposition**



CM-101 interferes with the core pathways that drive PSC

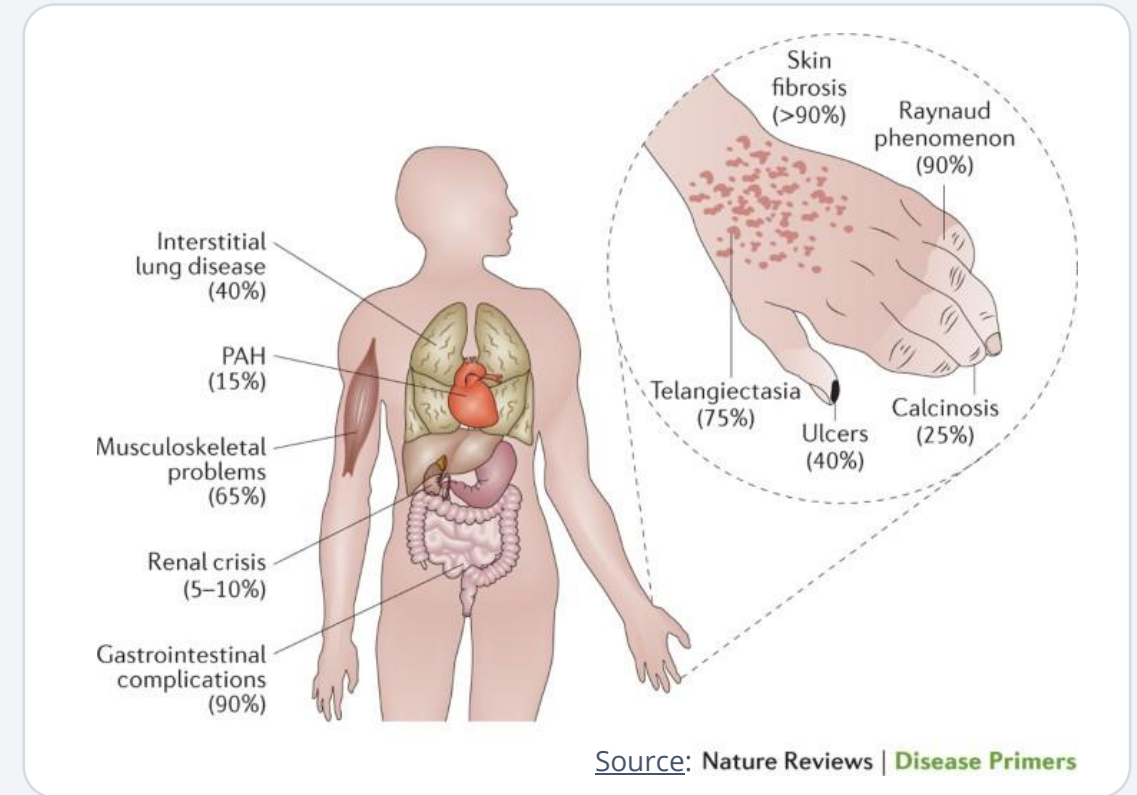
CM-101 for the Treatment of Systemic Sclerosis



Systemic Sclerosis (SSc)

Most Lethal Among Systemic Rheumatic Diseases

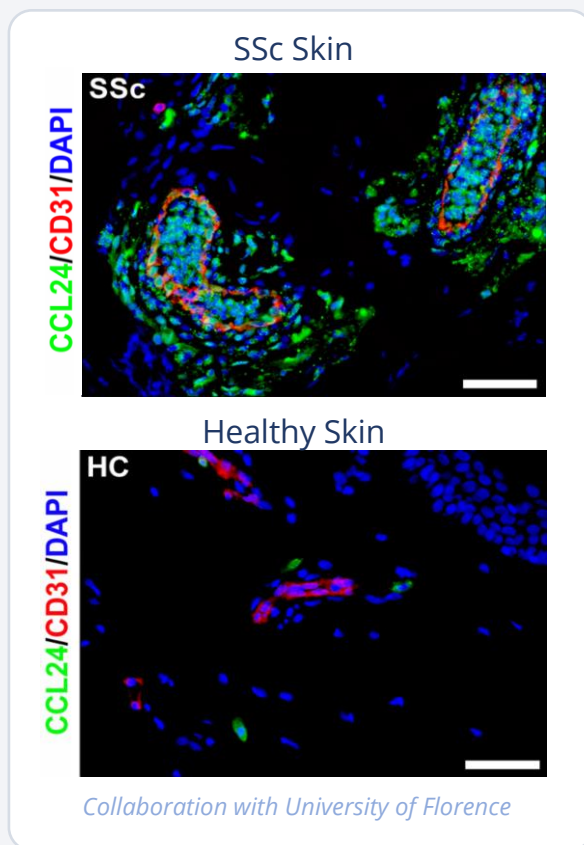
- **Rare** autoimmune rheumatic disease characterized by **inflammation and fibrosis** of the internal organs as well as vasculopathy
- **Age of diagnosis 30-50 years**, Predominantly affects women (3:1 female:male ratio)
- There is **no approved disease modifying drug**; Treatments focus on managing disease symptoms (Nintedanib, Tocilizumab)
- Median survival of 10 years - the **highest mortality rate among the systemic rheumatic diseases**
- Pulmonary involvement is the leading cause of death
- ~170K patients in 7 major markets



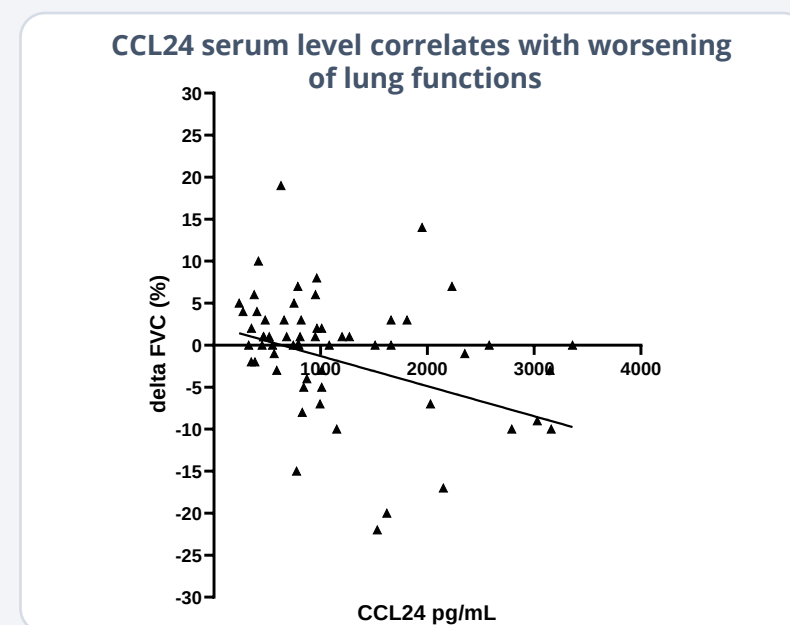
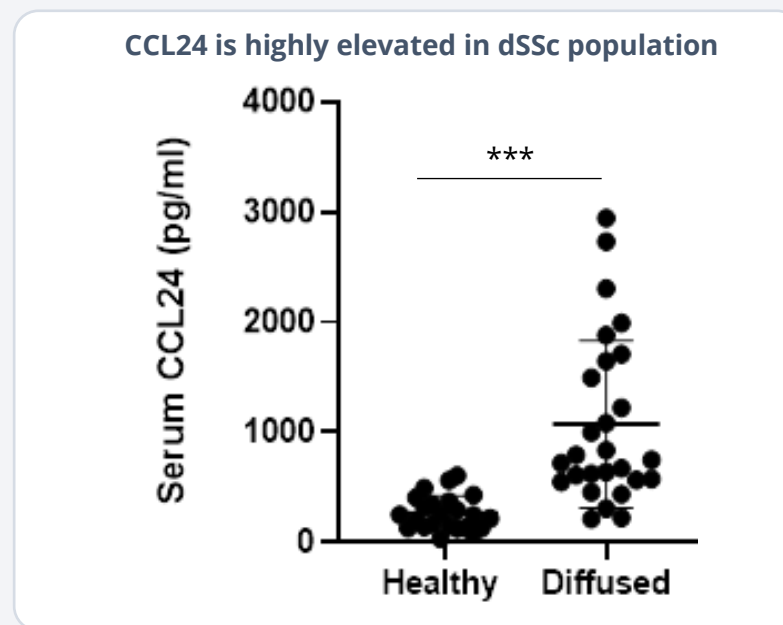
CCL24: A Critical Node Potentiating Systemic Sclerosis (SSc)



Clinical Relevance of CCL24 in SSc Pathophysiology is Supported by Patient Samples



CCL24 overexpressed in skin and serum samples of dSSc patients compared to healthy
CCL24 correlates with disease progression and fibrotic biomarkers

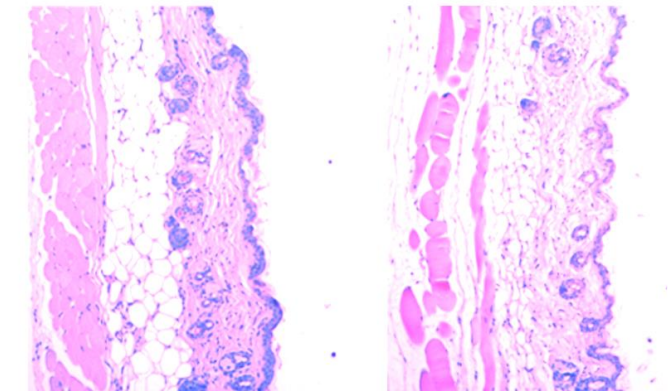
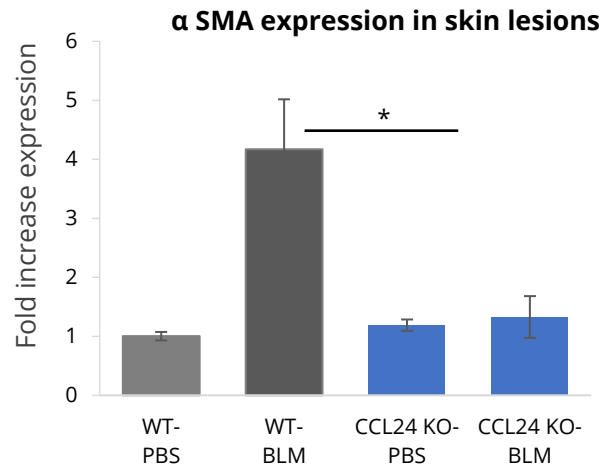
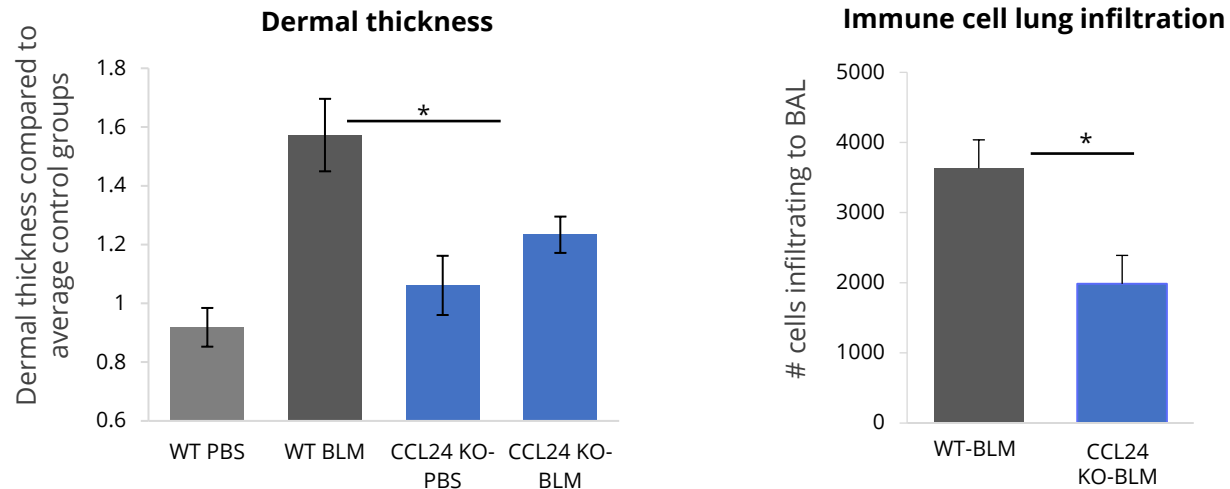


Patient samples support the targeted patient population and clinical trial endpoints

CCL24 Blockade is Singularly Sufficient to Attenuate Fibrosis

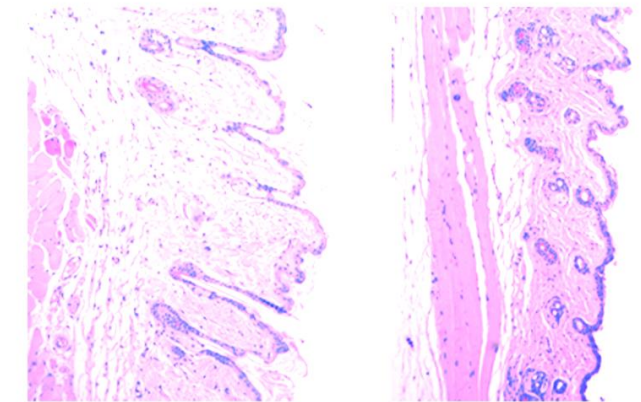


Knocking Out the CCL24 Gene Attenuates Inflammation and Fibrosis in SSc Bleomycin Experimental Model



CCL24 KO-BLM

CCL24 KO-PBS



WT-BLM

WT-PBS

Mor A et al., Annals of Rheumatic Diseases, 2019

BLM – Bleomycin; WT - Wild type; KO – Knockout, BAL- Bronchoalveolar lavage, PBS- Phosphate-buffered saline

*p $\leq$ 0.05

CM-101 Profoundly Reduces Skin and Lung Fibrosis in SSc

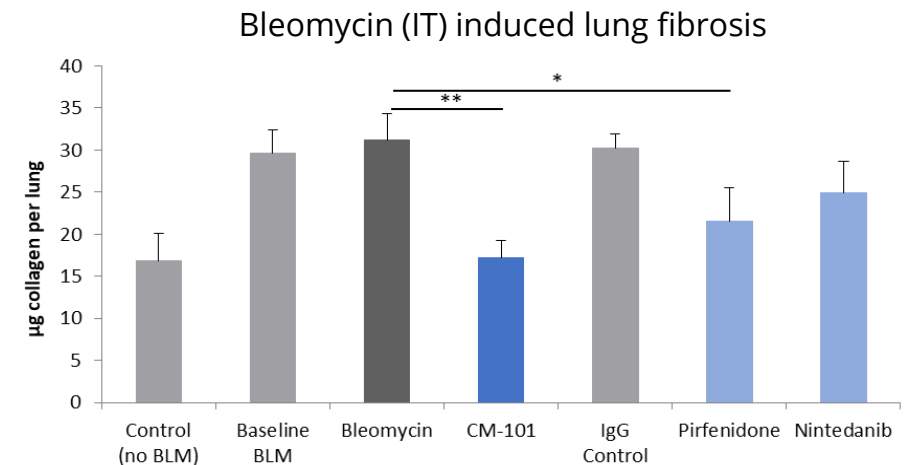
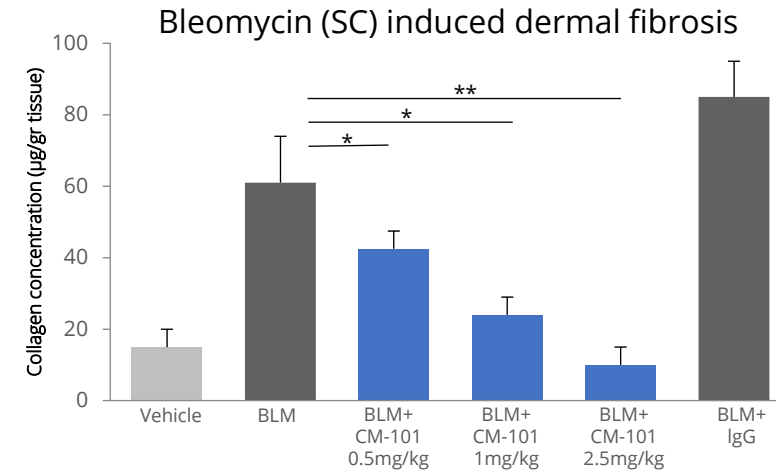
Experimental Models Relevant to Systemic Sclerosis Using Prevention and Therapeutic Designs



Attenuates skin and lung fibrosis levels in bleomycin-induced models using treatment mode

Demonstrates a **dose dependent attenuation of fibrosis**

Reduces lung collagen and inflammation compared to approved drugs for lung fibrosis



Mor et al, Annals of Rheumatoid Diseases, 2019

**p ≤ 0.01; *p ≤ 0.05.

BLM - bleomycin; IT - Intratracheal; SC - Subcutaneous

IgG - Immunoglobulin G

CM-101 Clinical Development Plan and Key Catalysts



Will be Advancing in Parallel Three Phase 2 Clinical Programs

	Status	2021	2022	2023
Primary Sclerosing Cholangitis	Enrolling Top line data expected in 2H2022	Phase 2: 15 wk treatment *Randomized, placebo-controlled; <i>US, Europe and Israel study sites</i>		
Systemic Sclerosis	Initiation expected in 1Q2022		Phase 2: 48 wk treatment Randomized, placebo-controlled; <i>US, Europe & Israel study sites</i>	
Liver Fibrosis Proof of MoA using SC formulation	Enrolling Top line data expected in 1H2022	Phase 2a: 16 wk treatment Randomized, placebo-controlled; <i>Israel study sites</i>		

SC – Subcutaneous, MoA- Mechanism of Action

*[NCT04595825](https://www.clinicaltrials.gov/study/NCT04595825)

Investment Summary

Pioneering Innovative Treatment for Fibrosis-Related Diseases with High Unmet Need



Chemomab

- Clinical stage company advancing Phase 2 trials in multiple fibrotic indications with high unmet need
- Substantial near term value inflection points
- Strong leadership with proven track record

CM-101

- First-in-class mAb blocking CCL24
- Novel and differentiated dual anti-fibrotic and anti-inflammatory MoA
- SC and IV formulations available; well established manufacturing process
- Strong IP protection

Efficacy

- First anti-fibrotic evidence in patients
- Significant anti-fibrotic effects across multiple *in vivo*, *ex vivo* and *in vitro* models

Safety

- Favorable safety and tolerability that support chronic treatment based on toxicology, Phase 1a and Phase 1b clinical trials

PK & Mode of Administration

- Optimal PK for both SC and IV formulations
- Comparable exposure levels and target engagement using both formulations

Thank You

