



# Pioneering Innovative Treatments for Fibrotic Diseases

Corporate Overview | Non-Confidential | March 2021

# Forward Looking Statements



This presentation contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act. These forward-looking statements include, among other things, statements regarding the clinical development pathway for CM-101; the proposed merger between Chemomab and Anchiano; expectations regarding ownership structure of the combined company; the future operations of the combined company and its ability to successfully initiate and complete clinical trials and achieve regulatory milestones; the nature, strategy and focus of the combined company; the development and commercial potential and potential benefits of any product candidates of the combined company; that the proposed merger will close and will enable the combined company to participate in the possible success of the combined company's product candidates; and that the product candidates have the potential to address high unmet needs of patients with serious fibrosis-related diseases and conditions. Any statements contained in this communication that are not statements of historical fact may be deemed to be forward-looking statements. These forward-looking statements are based upon Anchiano's and Chemomab's current expectations. Forward-looking statements involve risks and uncertainties.

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# Chemomab Highlights

*A Clinical Stage Biotech Company*



## Focus

- Development of innovative therapies for rare fibrotic diseases

## Clinical Differentiation

- CM-101, a first-in-class CCL24 neutralizing mAb with confirmed anti-fibrotic MoA
- Validated CCL24 as critical fibrosis target: clinical findings and experimental models
- Positive Ph1b data including safety, tolerability, PK, PD and biomarker readouts

## Near-Term Catalysts

- Initiating staggered phase 2 clinical programs, with first started in 4Q20
- Clinical readouts are expected during 2021-2022 to drive multiple value inflections

## Robust IP Portfolio

- Issued CoM, multiple nationalization stage filings, worldwide patent exclusivity through 2041

## Top Tier Investors



**THIEL**



# Experienced Leadership



## Management



**ADI MOR, PhD**

Chief Executive Officer, Co-Founder



**ARNON AHARON, MD**

Chief Medical Officer



**SIGAL FATTAL, CPA**

Chief Financial Officer



**SHARON ELKOBI, MSc, MBA**

VP Business Development



**MICHAL SEGAL-SALTO, PhD**

VP Research and Development



**SHARON HASHMUELI, PhD**

Head of CMC and Regulatory Affairs



## Board of Directors

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Chairman of the Board



**NISSIM DARVISH, MD, PhD**

Director



**JASON CAMM**

Director

**THIEL**

**ADI MOR, PhD**

Chief Executive Officer & Co-Founder



**DAVID BEN AMI**

Director



## Scientific Advisory Board

**Prof. Marco Matucci, Cerinic, MD, PhD**

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

**Prof. Dinesh Khanna MD, MBBS, MSc**

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

**Prof. Francesco Del Galdo, MD, PhD**

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

**Scott L. Friedman, MD**

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

**Massimo Pinzani, MD, PhD, FRCP**

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

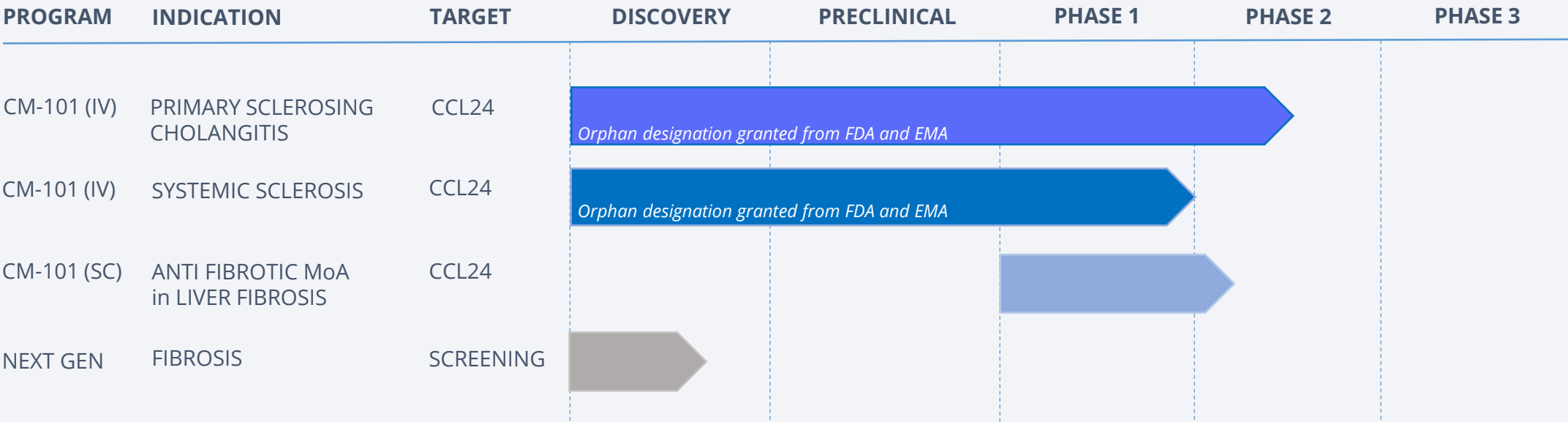
**Gideon Hirschfield, MA MB PhD**

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

# Chemomab Strategy: Fibrotic Disease Franchise



*Addressing Fibrotic Diseases with High Unmet Need*



# Fibrotic Diseases – Unmet Medical Need

*Potentiating High Morbidity and Mortality*

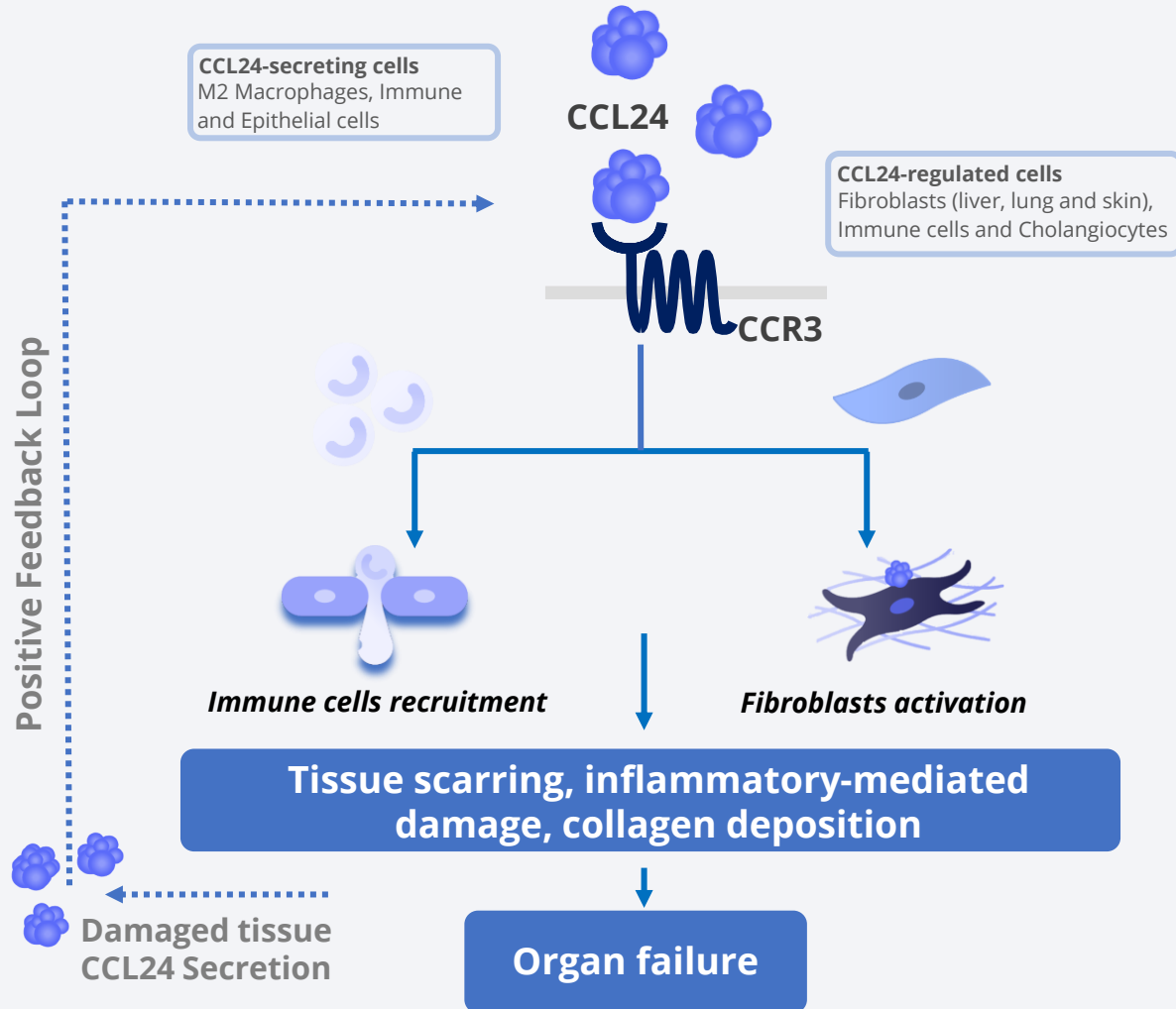


Fibrotic Disease	Orphan Disease	Sequela	SOC	Median Survival	7MM Prevalence	7MM Value
<b>Primary Sclerosing Cholangitis (PSC)</b>	✓	Bile duct damage, liver inflammation, fibrosis, cirrhosis, cholangiocarcinoma	Symptomatic approach, No FDA approved drug Liver Transplant - the only disease modifying option	10 – 12 yr	70K pts	>1B\$
<b>Systemic Sclerosis (SSc)</b>	✓	Skin fibrosis, vasculopathy multi-organ failure (lung, GI, heart)	Symptomatic approach, Nintedanib approved for ILD, No disease modifying option	10 yr	140K pts	>1B\$

No disease-modifying drugs have been approved for these fibrotic disorders

# CCL24 is a Novel Therapeutic Target for Fibrosis

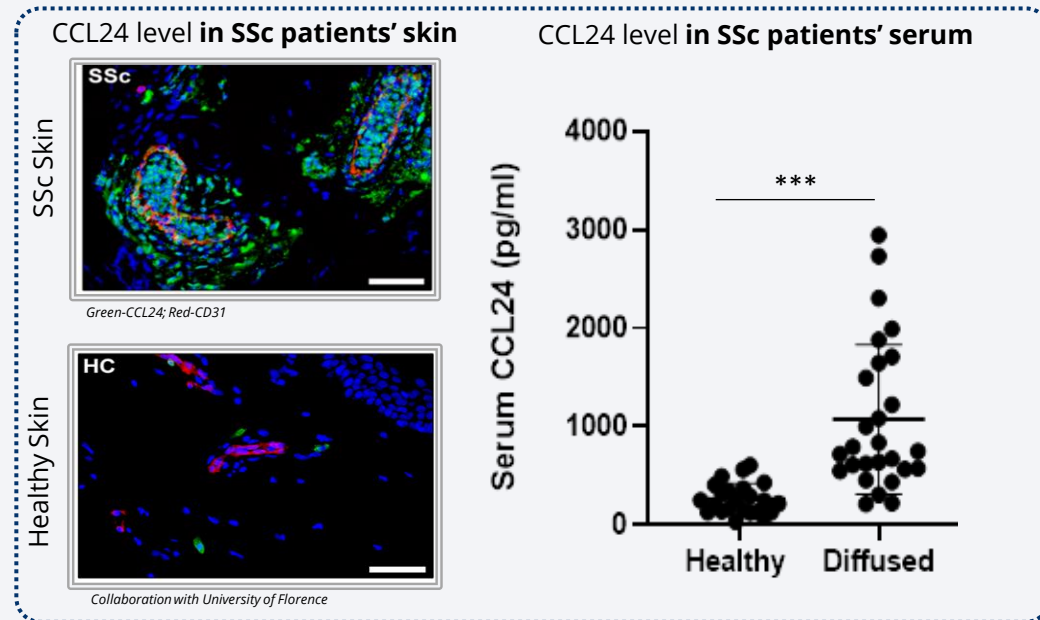
*Critical Mediator Promoting Inflammation and Fibrosis*



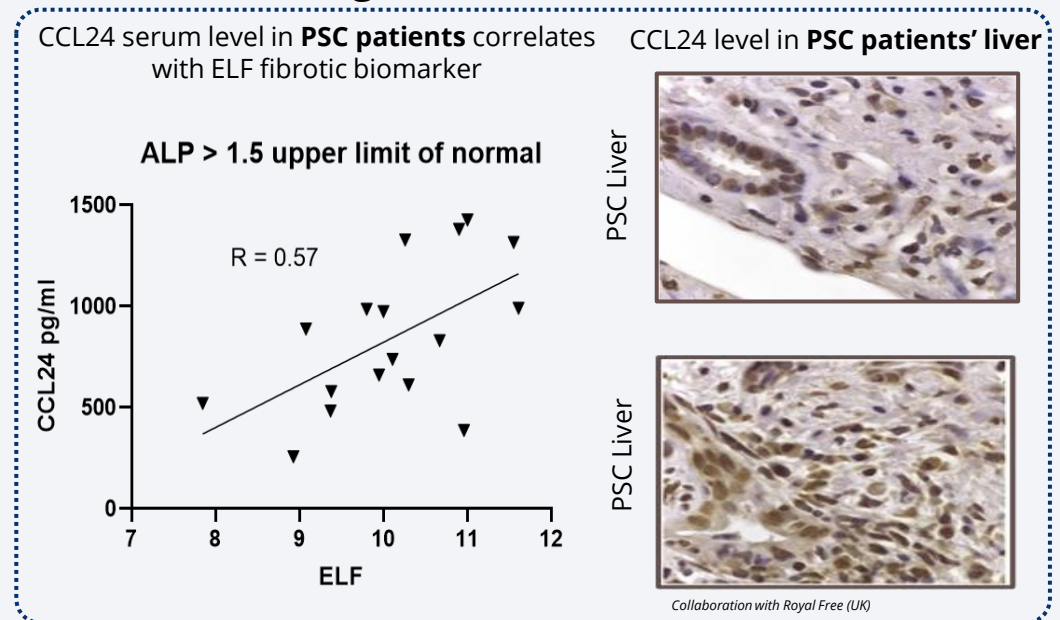
- ✓ **Dual role in promoting fibrosis**
  - directly activates fibroblasts
  - enhances local immune cell recruitment
- ✓ **Unique and differentiated activity**
  - ex vivo and in vivo data confirms unique role vs other CCLs
  - correlates with disease outcome and fibrotic biomarkers
- ✓ **Minor expression 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

# CCL24: a Critical Node Potentiating Fibrosis-Related Diseases

## Target Validation in SSc



## Target Validation in PSC

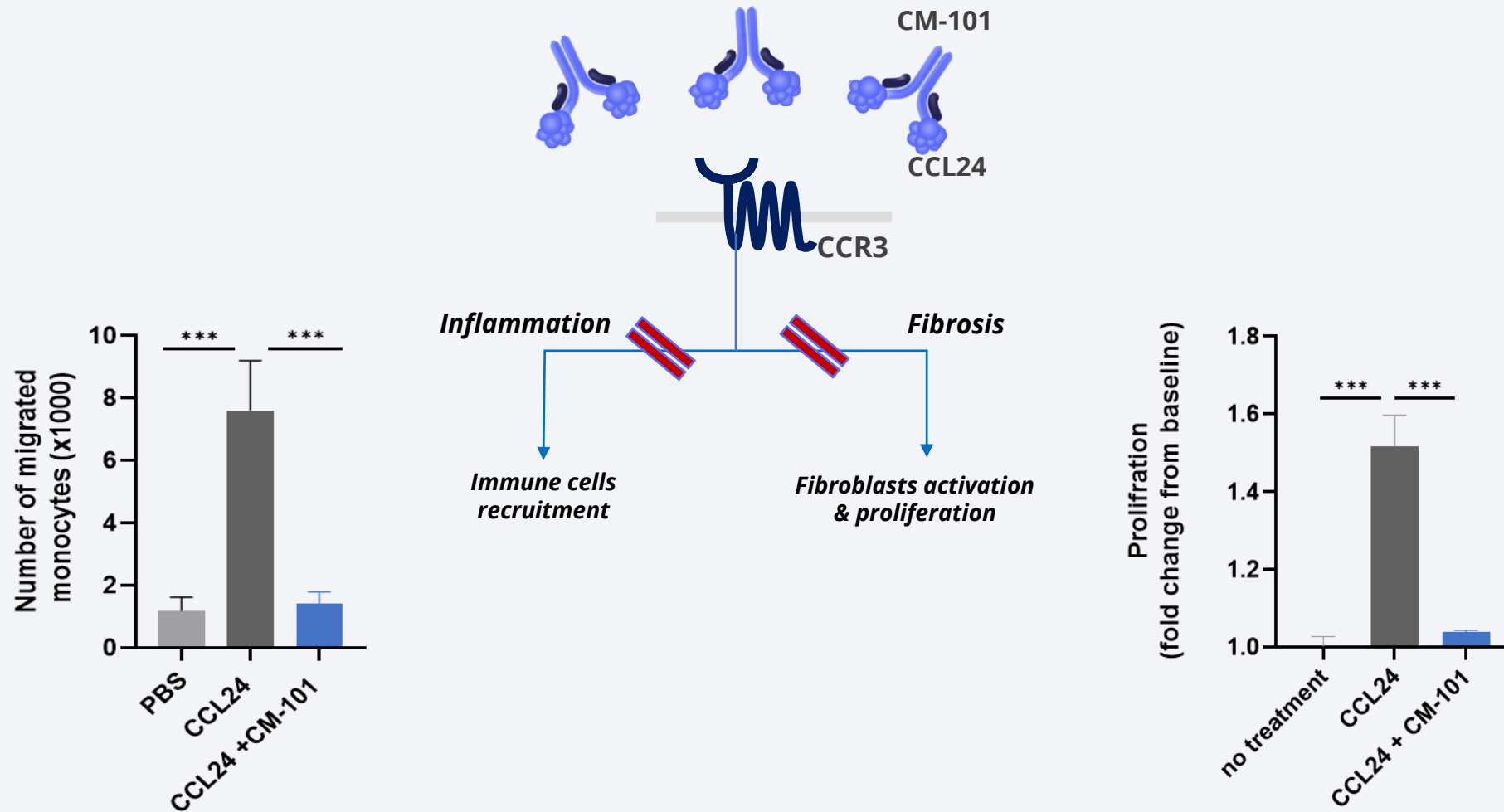


- CCL24 expression are **significantly and selectively elevated** in diseased vs. healthy tissues
- CCL24 levels **correlates with disease progression** and circulating fibrotic biomarkers
- CCL24 functionally **drives critical fibrotic** pathways across several experimental models
- CCL24 blockade using CM-101 demonstrates first evidence of **anti-fibrotic activity in human**



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

Dual Mechanism of Action Interfering with the Core Fibrotic Pathways

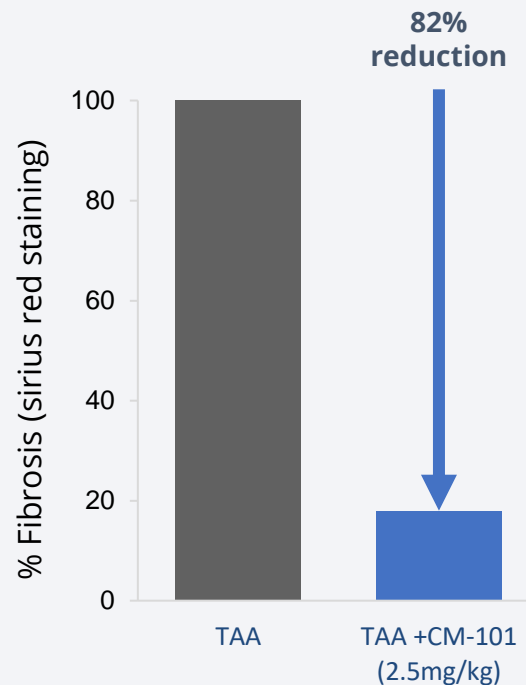
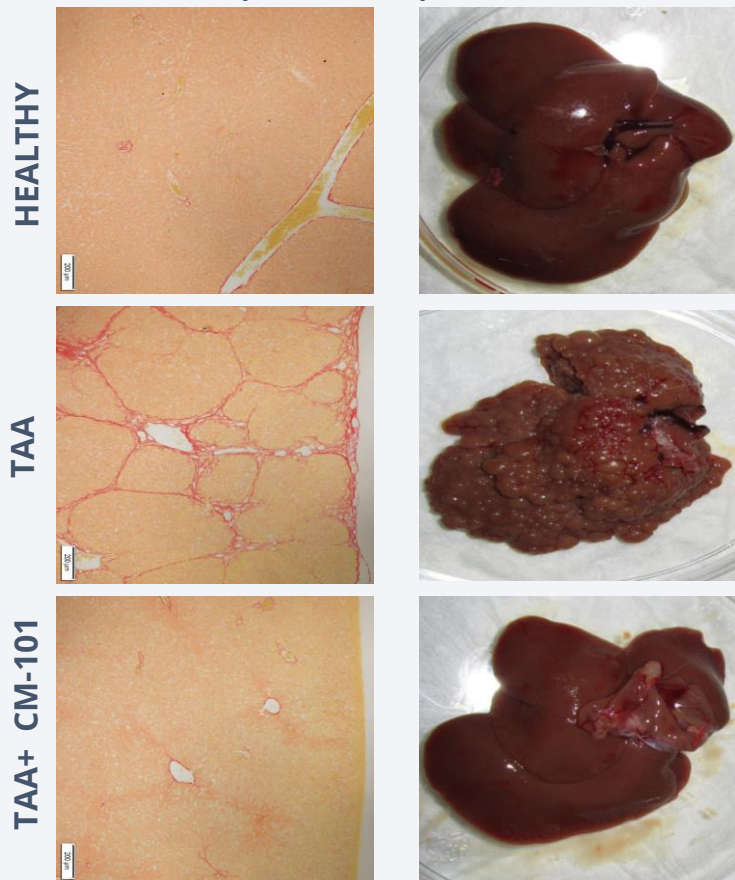


# CM-101 Reduces Liver Fibrosis by 80%

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



SIRIUS RED (COLLAGEN)



Pro-fibrotic genes

Col1A1  
Col3A1  
TIMP1  
ACTA2  
TGF- $\beta$

Liver enzymes

ALT  
AST  
ALP

Segal-Salto et al, JHEP reports 2020

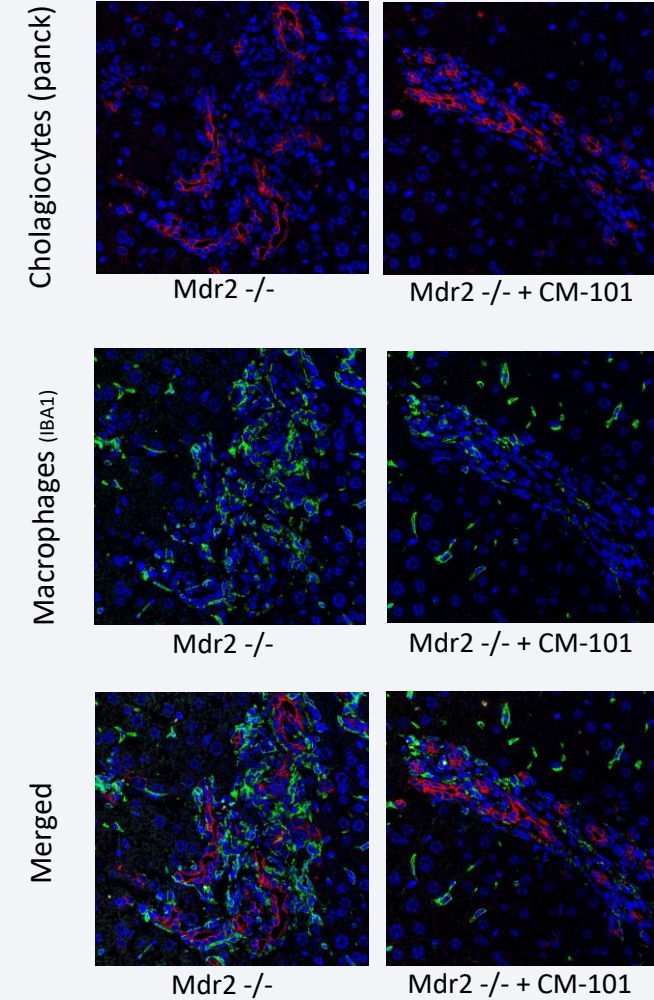
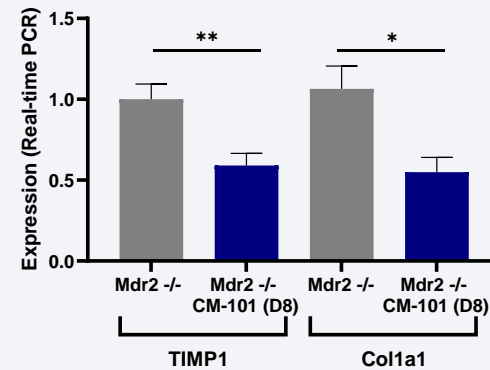
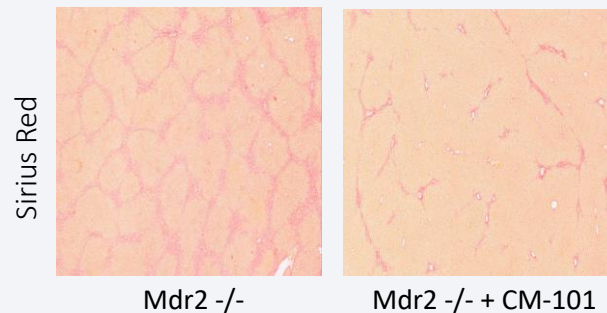
# CM-101 Reduces Liver Injury and Fibrosis in PSC

*Cholestasis, Inflammation and fibrosis are reduced in the MDR2 Knockout Model in Mice*



## CM-101 Interferes with the core mechanisms of PSC:

- Alkaline Phosphatase
- Bile acid
- Liver collagen levels
- Circulating and tissue resident Inflammatory cells
- Cholangiocytes proliferation



# CM-101 Profoundly Reduces Skin and Lung Fibrosis in SSc

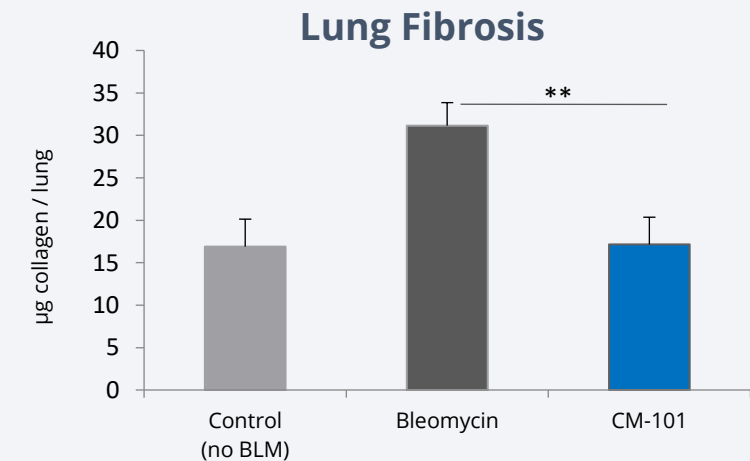
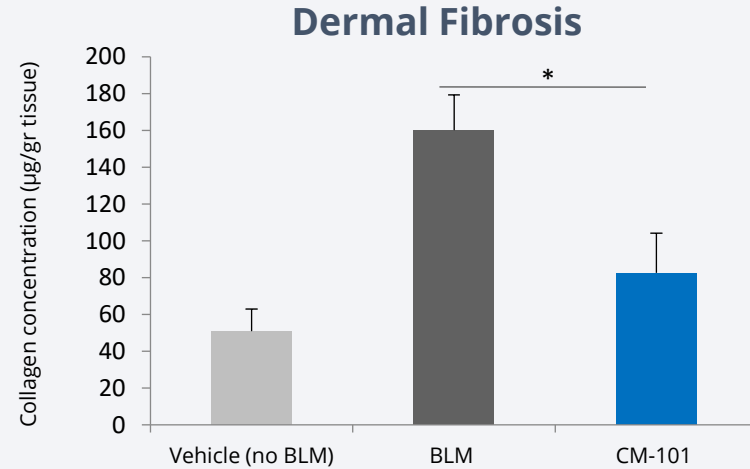
*Bleomycin induced SSc Models Using Therapeutic Design*



Attenuates skin and lung fibrosis levels using treatment mode

Demonstrates a dose dependent attenuation of fibrosis

Substantially reduces lung collagen and inflammation as compared to approved drugs for lung fibrosis



Mor et al, Annals of Rheumatoid Diseases, 2019

# CM-101 Holds a Robust Preclinical Package

*Significantly Attenuates Fibrosis & Inflammation Across a Wide Range of Models*



## CCL24 Target Validation

### Ex-Vivo (Patient Samples)

#### PSC

- Biomarkers correlation
- Overexpression of CCL24 and CCR3

#### Systemic Sclerosis

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

#### 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)

#### NASH

- CCL24 knock out 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)

### Nonalcoholic steatohepatitis

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

### Atherosclerosis

- ApoE knock out model (mice)

## Mechanism of Action

### CM-101 effects on fibroblasts activation

- Dermal, Hepatic and Lung fibroblast activation
- Dermal and liver fibroblast transition to myofibroblasts
- Hepatic fibroblast motility

### CM-101 effects on immune cells migration and recruitment

- Dermal fibroblast migration
- Monocyte polarization
- Monocytes recruitment

## Toxicology

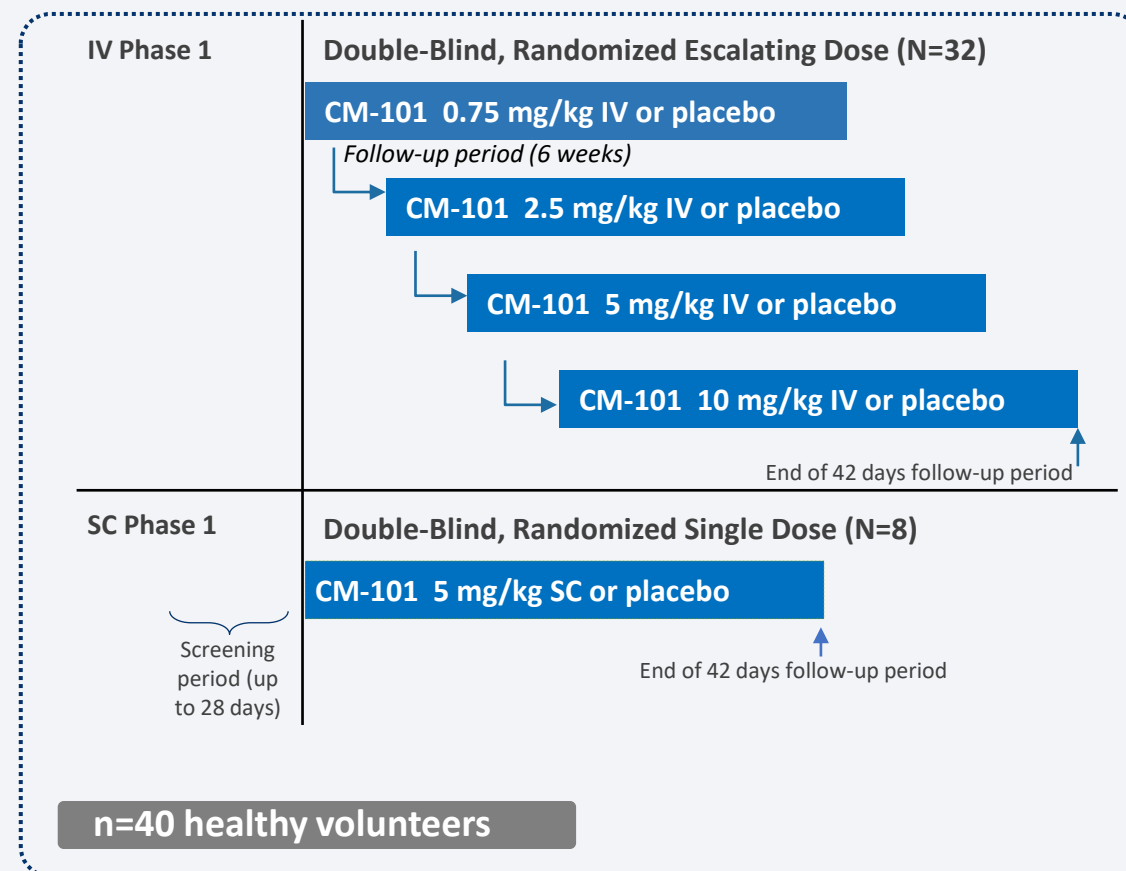
- **Short-term GLP** in rodents
- **Long-term GLP** in Non-human primates
- **Ex-vivo safety:** ADCC, CDC, cytokine secretion
- **Tissue cross reactivity**

# CM-101 is Safe & Well Tolerated in Healthy Volunteers

## Phase 1a Single Administration study



- CM-101 was **safe and well tolerated** at all tested doses up to 10 mg/kg and for both formulations
- Average  $t_{1/2}$  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

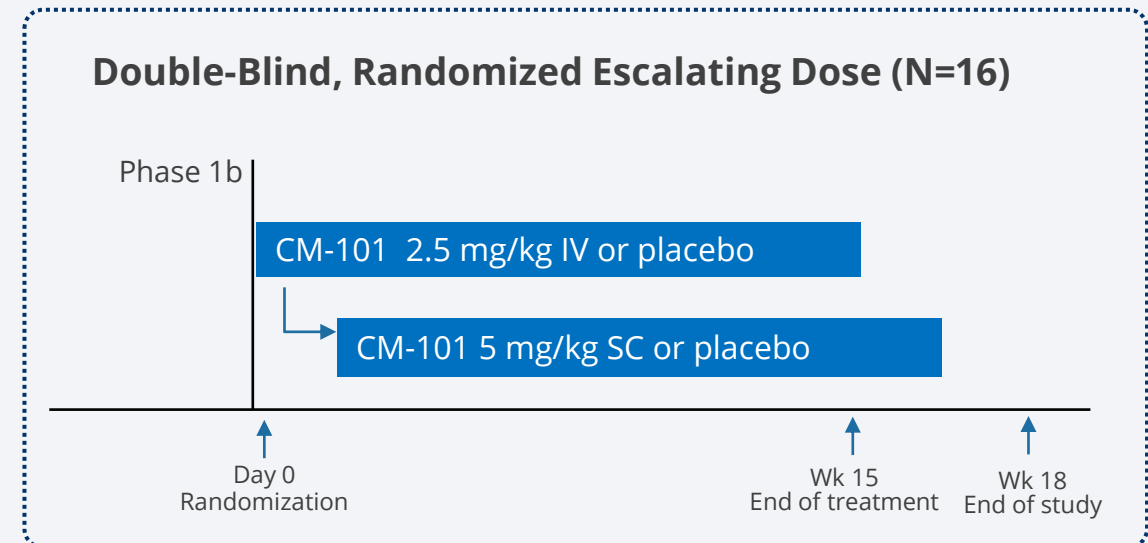


# Ph1b Demonstrates Safety & Tolerability Along 15 Weeks Treatment

Phase 1b Multiple Administration Study 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
- Favorable  $t_{1/2}$ , supports **long dosing interval** (Q2W - Q4W)
- Dose dependent **PK and target engagement**



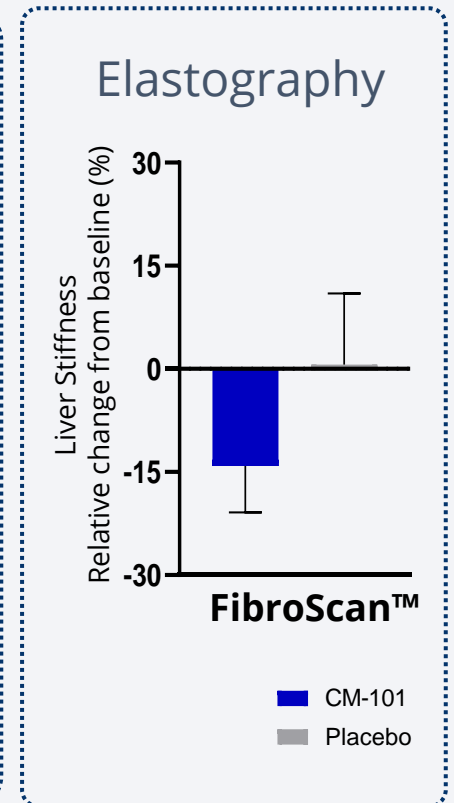
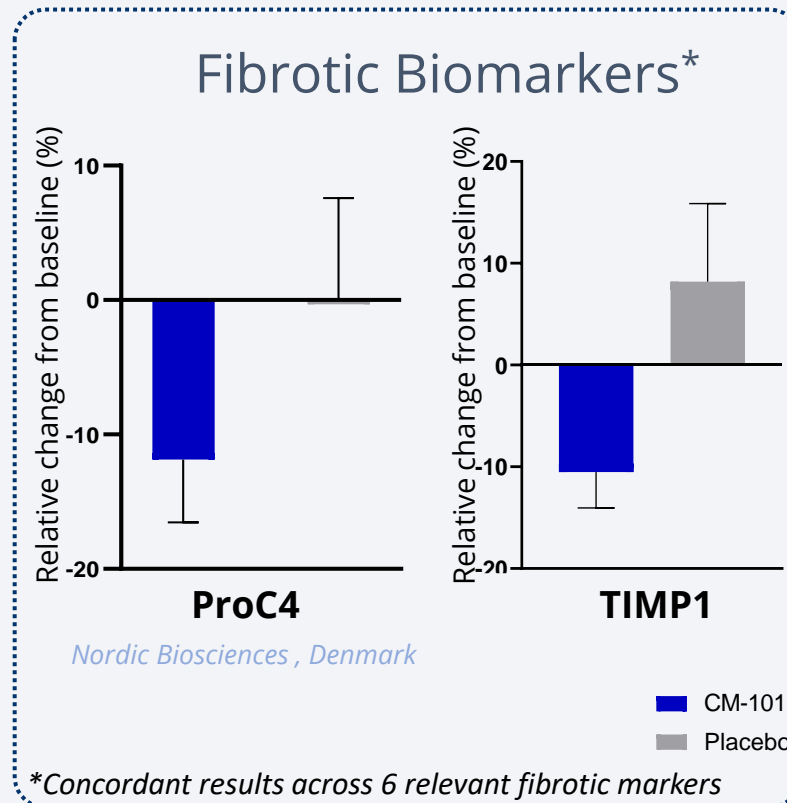
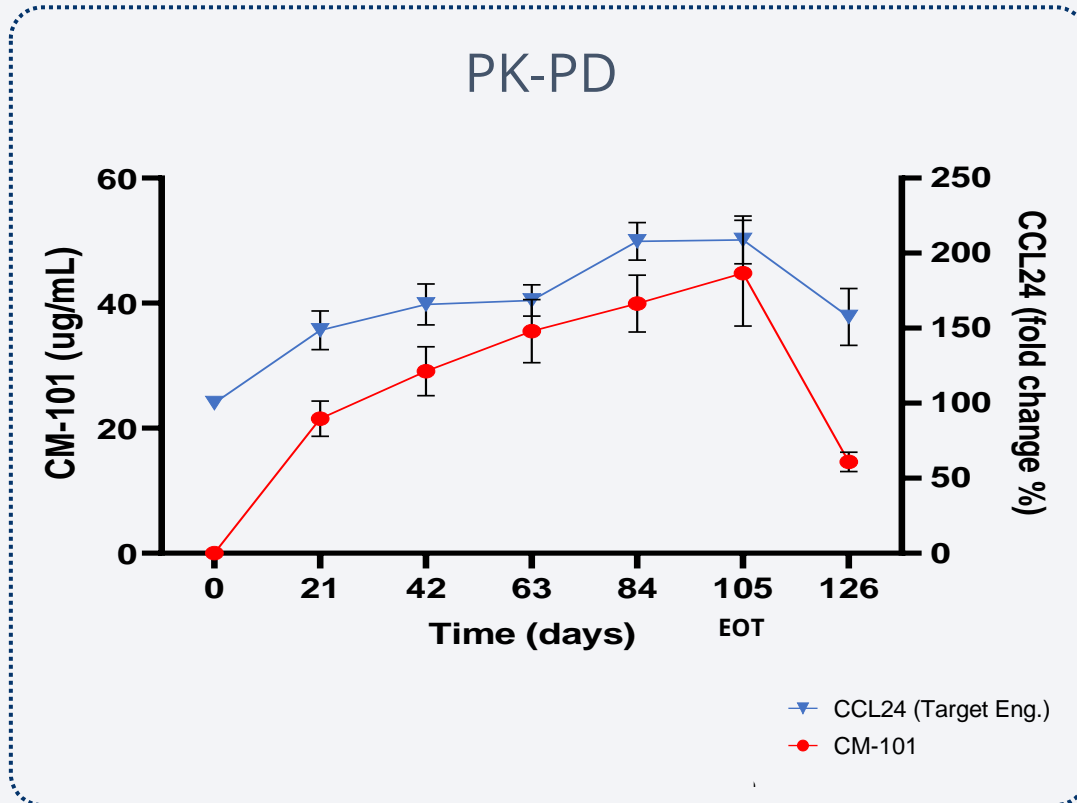
## Study Design

- 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

# CM-101 Target Engagement & Anti-Fibrotic Mechanism



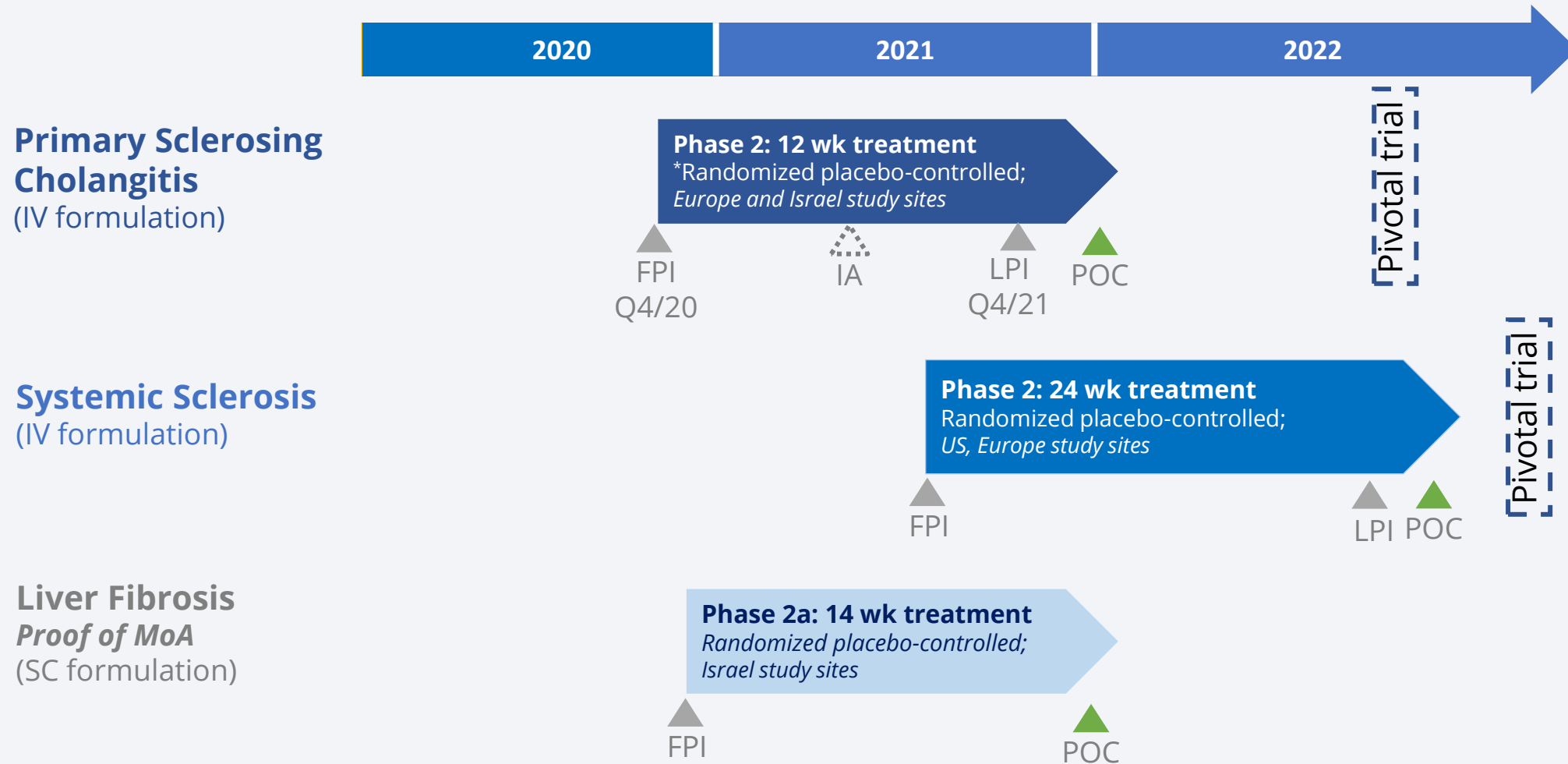
Human Confirmation for CM-101 Anti-Fibrotic Mechanism of Action





# CM-101 Clinical Development Plan and Key Catalysts

Phase 2 Studies Driving Pivotal Studies



# Chemomab, Fighting Fibrosis Across Indications



## Opportunity

- **Clinical stage company entering Ph2 trials in multiple fibrotic indications with high unmet need**
- **Substantial value inflection points in 2021-2022**
- **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 Formulation
- 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 Ia and Phase Ib 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

