

REPAIR. RESTORE. RENEW.™



The Wnt Company – Targeted Regeneration

May 2022

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From Wnt Gene Discovery to the Clinic

Scientific Discovery

1st **Wnt** gene
discovered
(Roel Nusse, Harold
Varmus)

1982

Biologic Validation

Surrozen
founded by The
Column Group in
collaboration with
preeminent Wnt
biologists

2016

2013

**Breakthrough
Prize in Life
Sciences** awarded
to Hans Clevers for
"describing the
role of Wnt
signaling in tissue
stem cells"

2017

**Breakthrough
Prize in Life
Sciences** awarded
to Roel Nusse for
"pioneering
research on the
Wnt pathway"

Therapeutic Transformation

First Wnt
modulating
antibody approved,
Amgen's Evenity
(romosuzumab) for
osteoporosis

2019

2020

Publication of
Surrozen's **SWAP**
and SWEETS
antibody platform
discoveries

Surrozen **progresses**
targeted Wnt
therapeutics
platform; expect to
initiate FIH trials

2022+

What is Wnt Biology?

Wnt Signaling Essential to Many Cell and Tissue Types

Fundamental Signal Transduction Biology

Wnt pathway central to:

- Regulating stem cell renewal, proliferation & differentiation
- Regenerating tissue

Wnt proteins generate array of Wnt signaling critical for:

- Shaping tissues during development
- Maintaining tissue architecture
- Repairing injured tissue

Many Organs and Tissues Require Wnt signaling



Eye



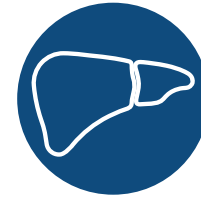
BBB



Lung



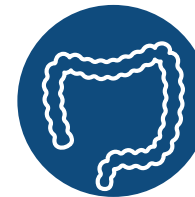
Cochlea



Liver



Pancreas

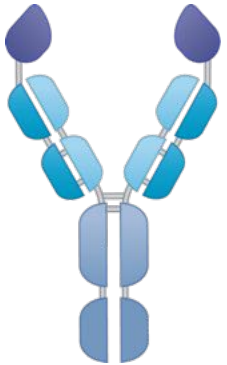


GI



Kidney

Surrozen – Leaders in Wnt Biology



Vision

Selectively target Wnt pathway to harness the body's own repair mechanism

Initial focus

Wnt related severe or acute diseases: GI, Liver, Ophthalmology

First in class

Proof of mechanism / biology and preclinical proof of safety for both lead candidates

Lead Product Candidates

Expect to initiate two clinical trials in Q3 '22

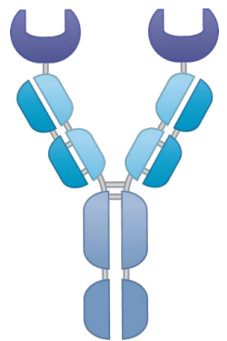
- SZN-1326: Inflammatory Bowel Disease
- SZN-043: Severe Liver Disease
- SZN-413: Ophthalmology

Proprietary platform

Unparalleled capabilities; demonstrated preclinical POC for several programs

Well positioned

\$104M cash balance



Our Novel Approach Overcomes Previous Challenges

Paving the Way to Targeted Antibody Regeneration

Potential first synthetic soluble Wnt mimetics

Selectivity: Target specific Fzd or cell surface receptors

Potency: Confer potency through multivalent binding

Safety: Mimic normal physiologic responses

Manufacturing: Easily manufacturable leveraging typical antibody methods with high yields

Validation of Our Prominent Role in Wnt Biology Breakthroughs

nature

Surrogate Wnt agonists that phenocopy canonical Wnt and β -catenin signaling

 CellPress



Development of Potent, Selective Surrogate Wnt Molecules and Their Application in Defining Frizzled Requirements

**SCIENTIFIC
REPORTS**
nature research

Tissue-targeted R-spondin mimetics for liver regeneration

Science

Structural Basis of Wnt Recognition by Frizzled

Fully Integrated, Repeatable Discovery Capabilities

Potential to Transform Patient Outcomes

Internal Capabilities

Wnt Biology Expertise

**Wnt Modulating
Antibody Engineering**

Wnt Pathway Profiling

Scientifically Driven Strategy

Scientifically Driven Strategy

~60 R&D employees

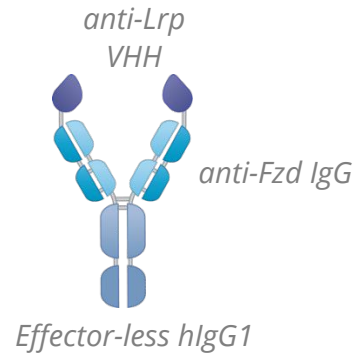
~ 50% PhD, MDs or PhD/MDs

Focus on diseases with compelling Wnt biology

Employ models with translatability to human disease

Proprietary Technologies Enable Selective Wnt Signaling

SWAP Technology

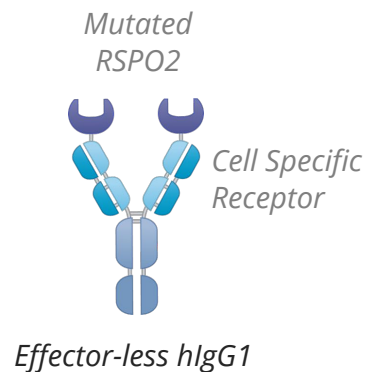


Mimic normal physiologic **response** (natural Wnt or natural R-spondin)

Applied in diseases with **deficient Wnt ligand** or **Wnt signaling**

Customized for each disease state

SWEETS Technology



Targeted with Fzd receptor selectivity or cell specific receptors

Deep Wnt Signaling Expertise Supports Productive & Expanding R&D Pipeline

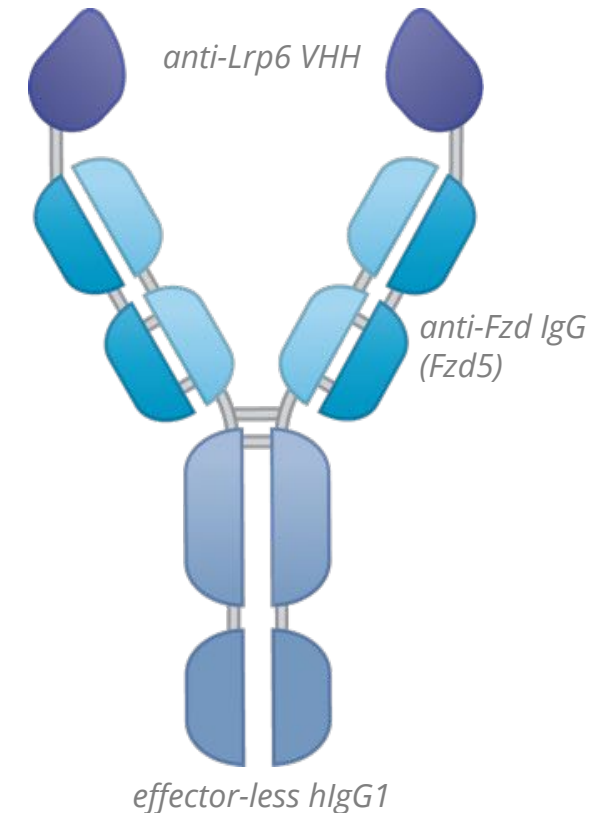
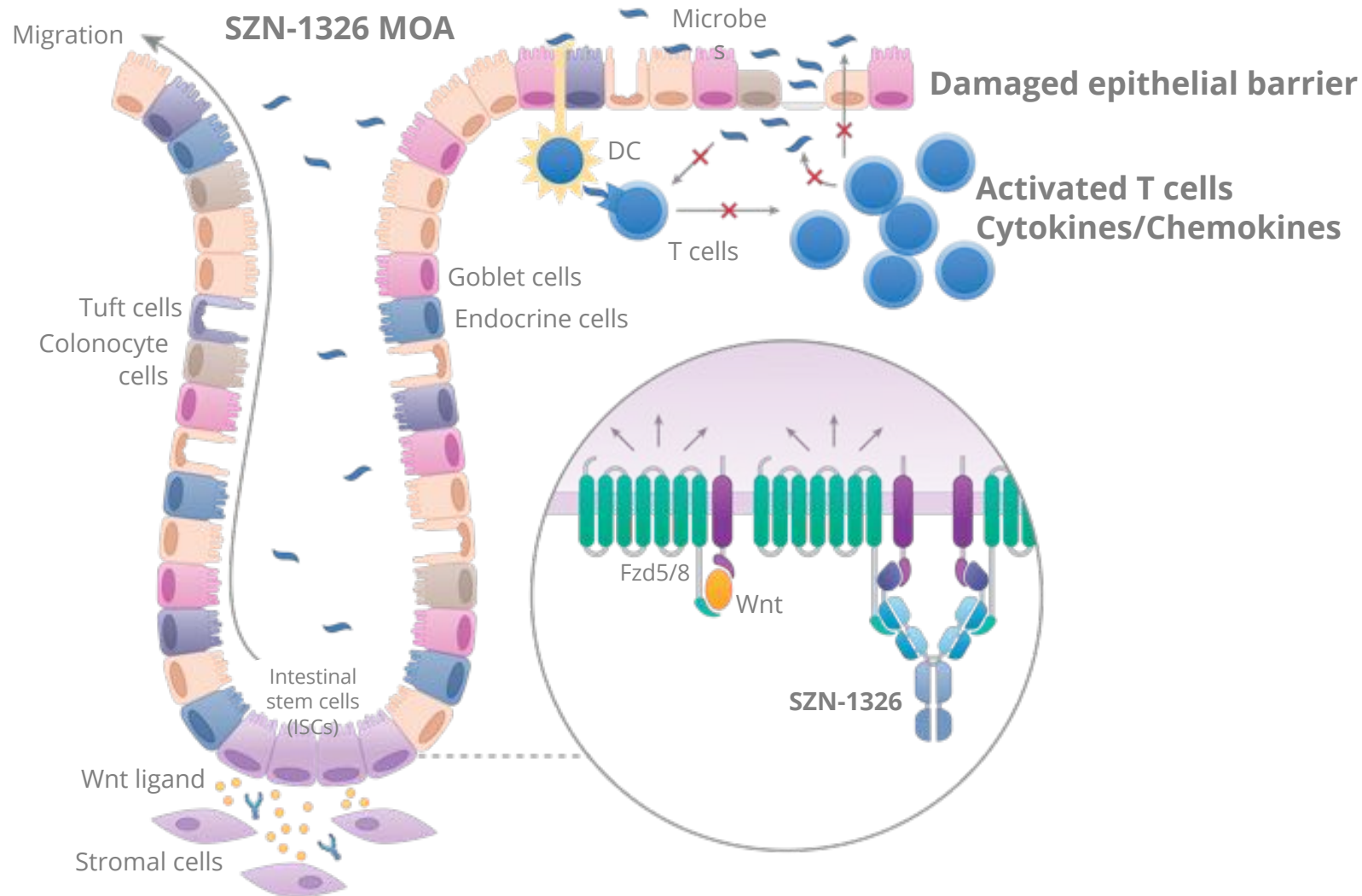
Lead Programs	Indication(s)	Research	Preclinical	Phase 1	Phase 2	Phase 3	Regulatory	Next Milestone
SZN-1326	Moderate to Severe IBD							Initiate clinical trial Q3'22
SZN-043	Severe Alcoholic Hepatitis							Initiate clinical trial Q3'22

Research Programs

Tissue	Indications	Discovery	Proof of Concept	Lead Candidate/s
Retinal Vasculature	Retinopathies			Nominated candidate Q1'22
Cornea	Fuchs' Dystrophy, Limbal Cell Def			
RPE	Dry AMD			
Lacrimal Gland	Severe Dry Eye (Sjögren's)			
Intestine	Short Bowel Syndrome			
Cochlea	Hearing Loss			
Lung	IPF, COPD			
Renal	Polycystic Kidney Disease, FSGS			

SZN-1326 – Intestine Targeted Epithelial Restoration

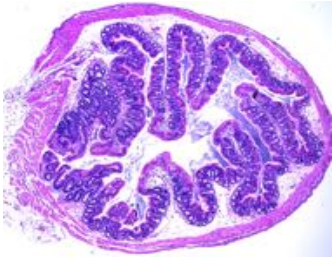
Mechanism Suggests Potential New Treatment Paradigm in Inflammatory Bowel Disease



SZN-1326 – Potential to Transform Treatment Paradigm in UC

Targeted antibody designed to repair epithelial barrier; induce mucosal healing

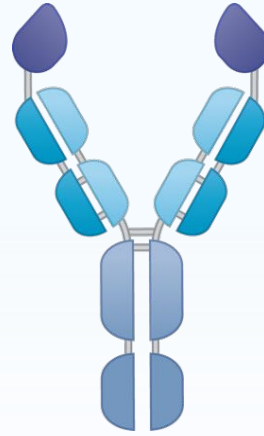
Background



Moderate to Severe Ulcerative Colitis

- Characterized by large intestine inflammation and ulcers
- Debilitating: frequent diarrhea, bloody stools, weight loss, dehydration, and anemia
- Complications from severe and chronic inflammation can become life-threatening
- SOC: Treated with anti-inflammatory agents
 - Takes months to induce remission
 - Achieve remission in < 50% and mucosal healing in < 20%
 - Fail multiple therapies

Our Solution



MOA: Designed to repair epithelial barrier & induce mucosal healing

- Dysregulation of Wnt signaling may play a role in abnormal epithelial healing in IBD
- Mucosal healing associated with better patient outcomes

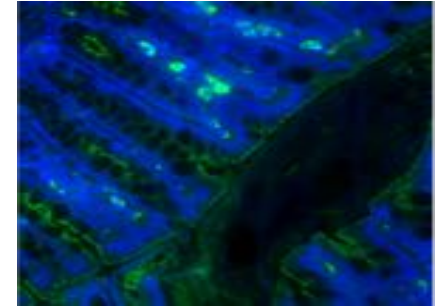
Targeted: Selectively targets Fzd5 abundant in intestinal epithelium

Intestine-Targeted Regeneration and Functional Improvement

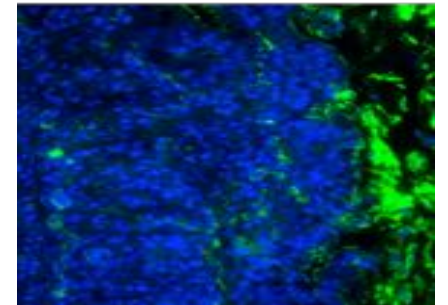
Differentiated Preclinical Data

- Repairs damaged colon epithelium
- Induces mucosal healing
- Reduces inflammation
- Improves disease activity index
- Better activity than other anti-inflammatory agents including biologics
- No adverse findings in GLP tox studies

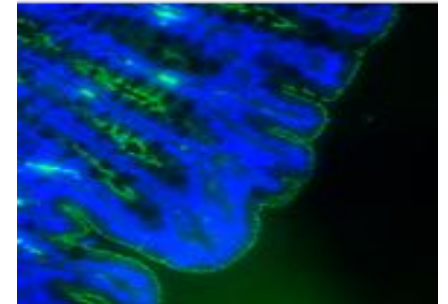
Normal
(No DSS Damage)



Damaged
(DSS Damage)



Restored
(DSS Damage + SZN-1326)



SZN-1326 Phase 1 Trial Overview

Focus - Proof of Clinical Concept in Ph 1b Ulcerative Colitis

Three-Part Ph 1 Randomized Trial Design

Ph 1a – SAD

Healthy volunteers

N = up to 44

Up to 5 randomized IV cohorts, and 2 SC cohorts

Ph 1a – MAD

Healthy volunteers

N = up to 24

Up to 3 randomized cohorts (IV)
Dosing for 4 doses

Ph 1b – MAD

Moderate-severe patients with UC

N= up to 24

Proof of clinical concept

Up to 3 randomized cohorts dosed IV weekly or biweekly for 12 wks. Follow-up - 24 weeks

Key Endpoints

Ph 1a SAD/MAD

- Safety
- PK/PD
- ADA

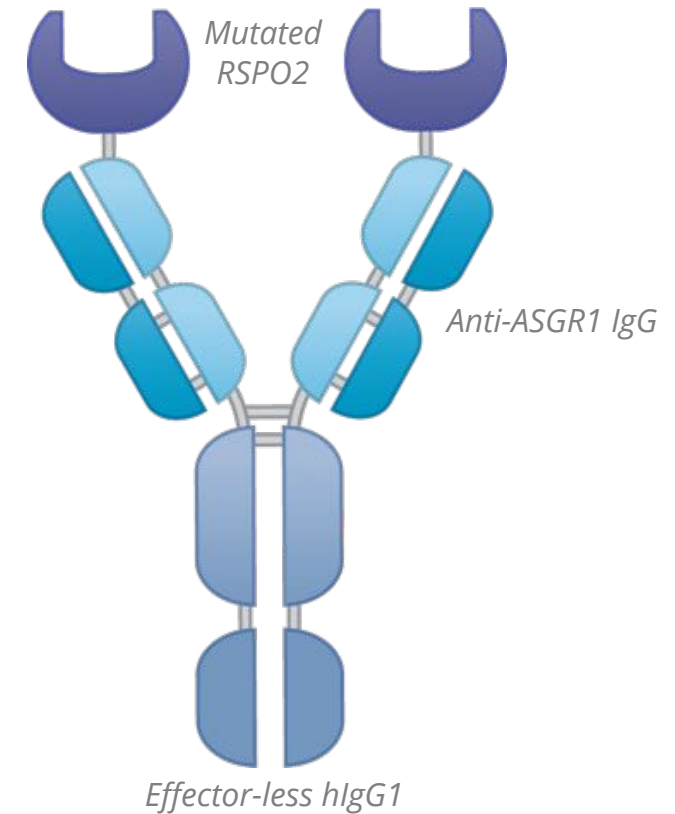
Ph 1b UC MAD

- Clinical remission and response
- Endoscopic remission
- Histologic remission
- UC-100
- PD markers

Potential for First Approved Treatment for Severe Alcoholic Hepatitis

Liver Specific Wnt Activation and Regeneration

SZN-043 MOA

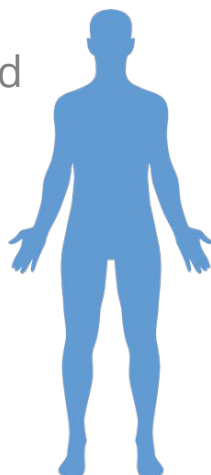


SZN-043 Potential to Transform Patient Outcomes in Severe AH

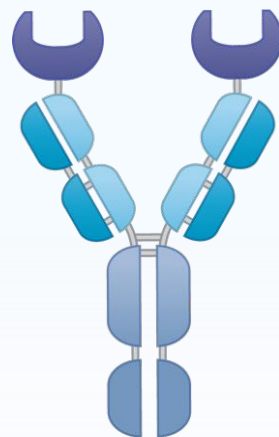
Targeted antibody designed to induce hepatocyte proliferation and improve liver function

Background

- Serious form of acute decompensated alcoholic liver disease caused by heavy alcohol use
- Leads to liver cell death, damage and subsequent inflammation
- 90-day mortality of 30%
- ~130K hospitalizations per year
- No approved treatments
 - Steroids: contra-indicated in > 50% of patients; no benefit at 3 months+
 - Liver transplants: limited supply, costly and often denied



Our Solution



MOA: SZN-043 designed to addresses underlying pathophysiology

- Hepatocyte proliferation correlated with increased survival
- Upregulation of Wnt signaling implicated in improved liver function

Targeted: Selectivity achieved through inclusion of ASGR1

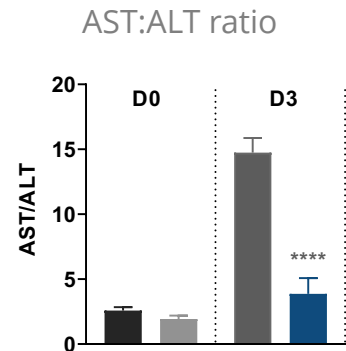
SZN-043 In Vivo Effects

Liver Specific Proliferation, Functional Improvement, Fibrosis Regression

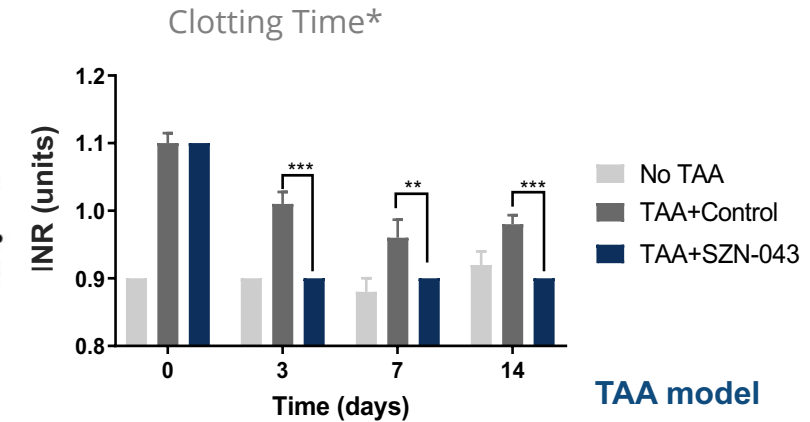
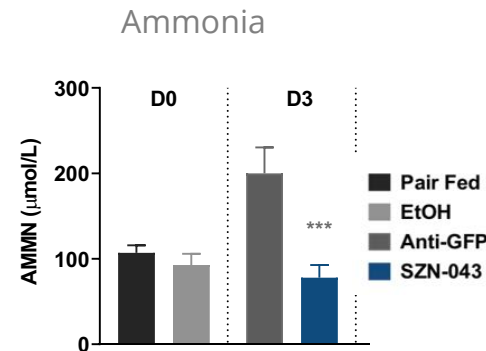
Compelling Preclinical Data

- >25 preclinical studies conducted
- Selectively activates Wnt Signaling
- Induces hepatocyte proliferation
- Rapidly improves liver function
- Reduces markers of liver injury & inflammation
- No adverse findings in GLP tox studies

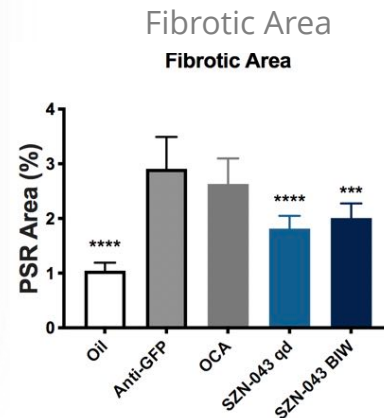
Improvement in Liver Function



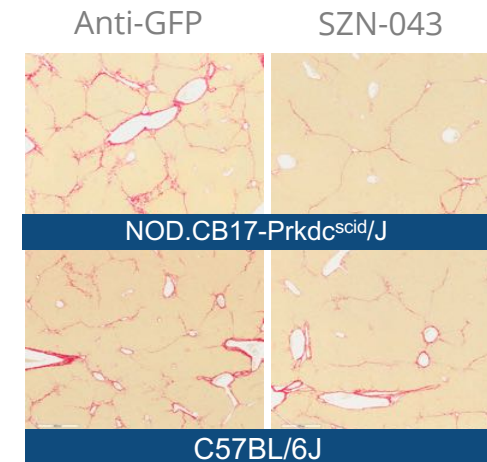
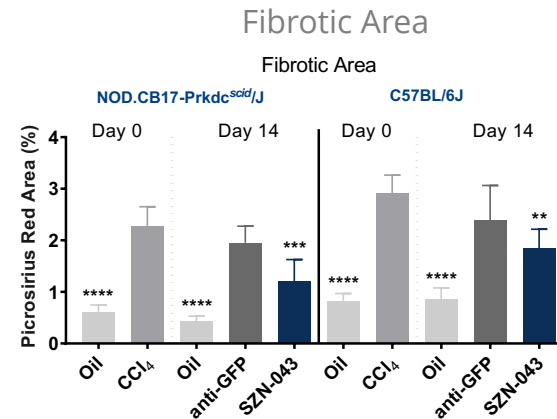
Alcohol liver injury model



Regression of Fibrosis



*SZN-043 V.2 in TAA Model



SZN-043 Phase 1 Clinical Trial Overview

Focus – Proof of Concept in Early Cirrhosis; Potential Expedited Regulatory Pathway

Multi-Part Ph 1 Randomized Trial Design

Ph 1a – SAD

Healthy volunteers

N = up to 24

Up to 4 randomized cohorts (IV)

Ph 1b – SAD/MAD

Early cirrhosis

N = Up to 16

Up to 2 randomized cohorts (IV)

PD markers indicative of liver proliferation and function improvement

Ph 1b

Severe Alcoholic Hepatitis (AH)

N = up to 30

Early read on LILLE score and MELD scores – high survival correlation

Further *proof of clinical activity*; potential for Fast Track and Breakthrough Designation



Key Endpoints

Ph 1a SAD:

- Safety, ADA
- PK/PD (including methacetin breath test)

Ph 1b SAD/ MAD - (early cirrhosis)

- Safety
- PK/PD (including methacetin breath test, Hepquant)
- ADA

Ph 1b Severe AH MAD

- Lille and MELD scores
- Mortality

Platform Broadly Applicable Across Wide Spectrum of Diseases

Current Focus

Current Technology



Gastrointestinal/ Liver

UC, Crohn's, SAH, ACLF



Ophthalmology

Wet AMD, Diabetic Retinopathy, DME, Severe Dry Eye (Sjögren's)

Current Focus

Future Technology

Future Opportunities

Current Technology

Kidney

FSGS, PKD



Lung

IPF, COPD



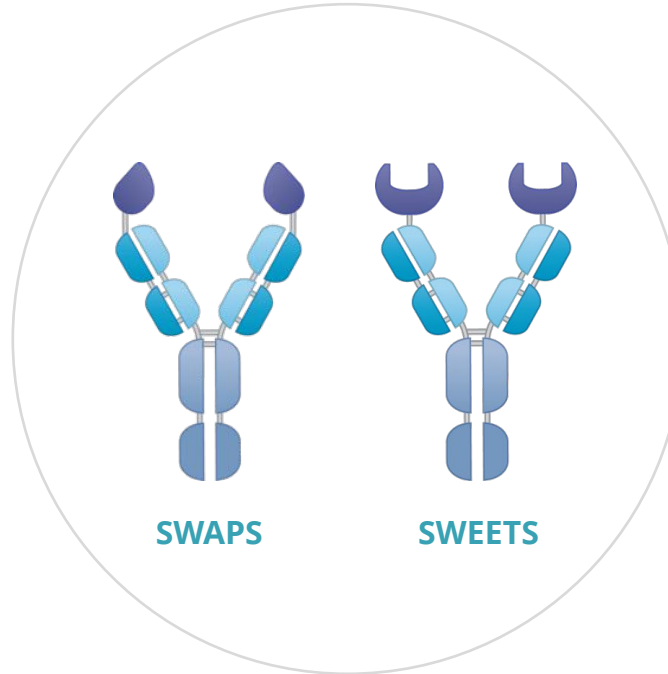
Neurology

MS, Stroke



Future Opportunities

Future Technology



Robust Activity in Multiple Preclinical Ophthalmology Models

SZN-413 (mono Fzd 4) lead candidate for retinopathy – addresses retinal non-perfusion and vascular leakage simultaneously

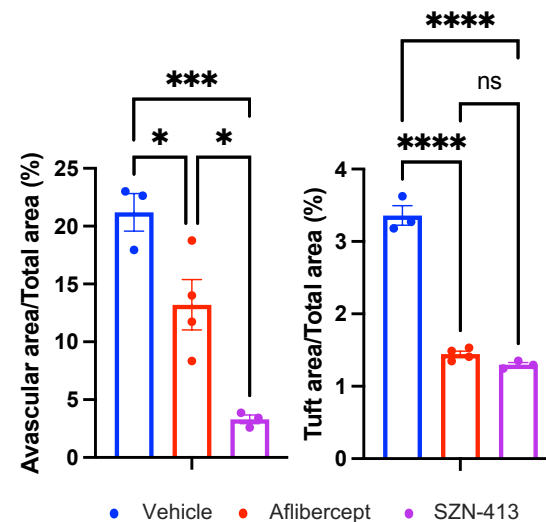
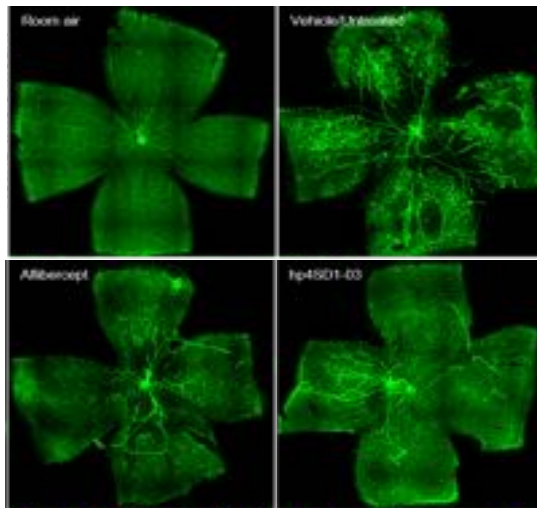
Fzd4 signaling plays critical role in retinal vasculature integrity

Stimulated Wnt signaling

Increased tight junction protein expression in endothelial cells

Restored norrin function in Ndp KO mice

Reduced avascular area & pathologic NV tuft formation in OIR model; reduced vascular leakage in VEGF-induced retinal model



Lacrimal Gland (LG) Program

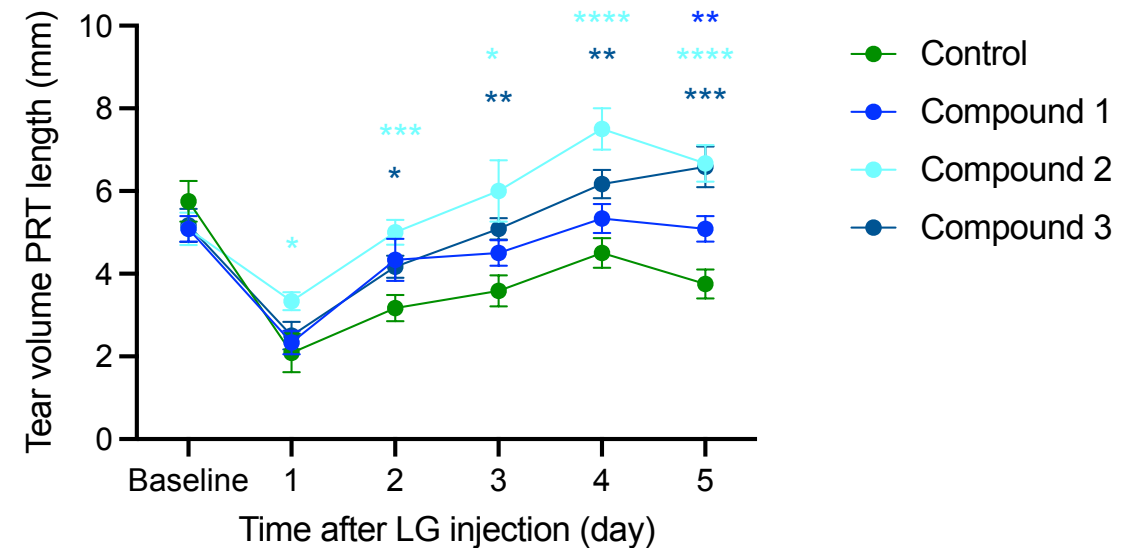
Tear-producing glands rely on Wnt signaling for function

Stimulated Wnt signaling

Effect observed in lacrimal and meibomian gland

Increased tear production within 2 days in IL-1a lacrimal gland model

Tear secreted by ipsilateral eye



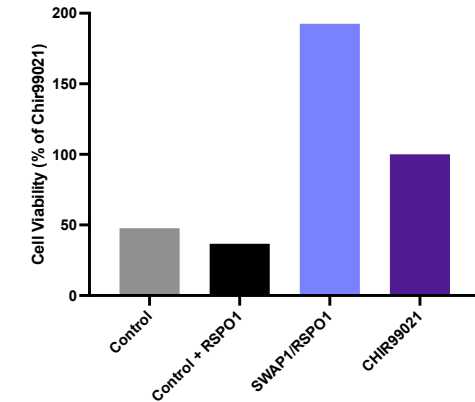
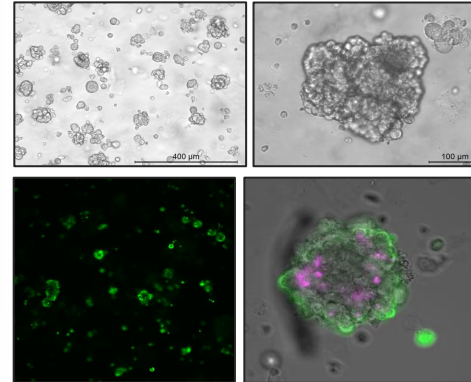
Lung Regeneration Program

Recent Discoveries Suggest Potential Role for Wnt in Treatment of IPF and COPD

Compelling Preclinical Data

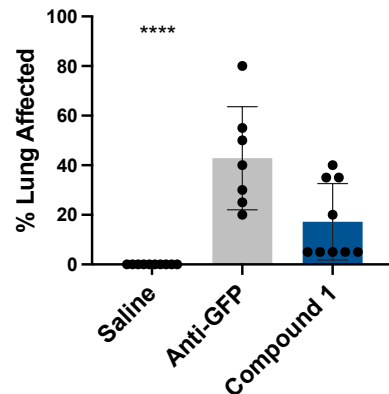
- Activates Wnt Signaling
- Expands alveolar AT2 cell organoids
- Reduced injury and improved fibrosis in the acute bleomycin model

Expands AT2 Cell Alveolar Organoids

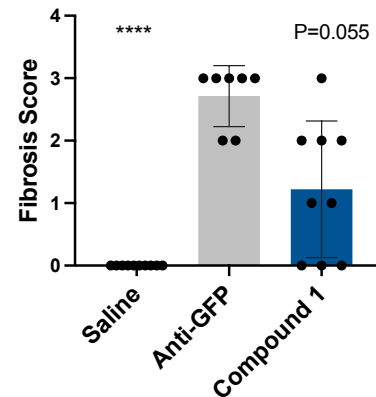


Reduction of Fibrosis

% Lung affected
(mice excluded with <5% BW change)

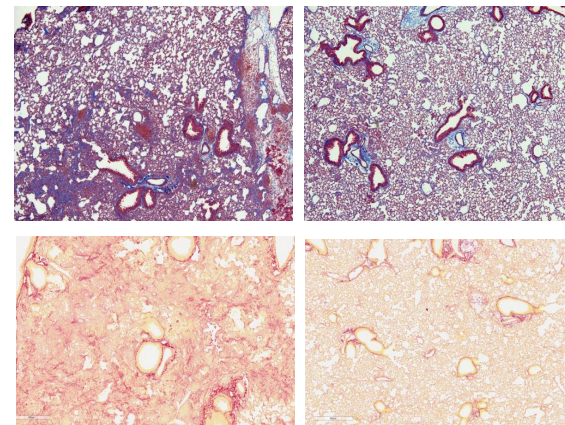


Fibrosis score
(mice excluded with <5% BW change)

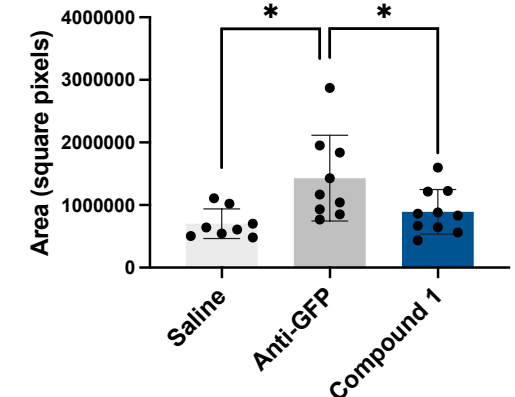


Anti-GFP

Surrozen



ACTA2 quantification



Near Term Outlook and Potential Milestones

Multiple Clinical Milestones with Potential for Early Proof of Concept

SZN-1326

Intestine

2021

Completed IND-enabling
Toxicology Studies

Q3' 2022

Phase 1a in Healthy Volunteers

2023

Phase 1b in Ulcerative Colitis
Patients

SZN-043

Liver

2021

Completed IND-enabling
Toxicology Studies

Q3' 2022

Phase 1a/b in Healthy Volunteers
and Early Cirrhosis Patients

2023

Phase 1b in Severe AH Patients

Research Programs

2022

Nominated Additional Lead
Candidate

2023+

Nominate Additional Lead
Candidate(s) and File INDs

REPAIR. RESTORE. RENEW.™



The Wnt Company – Targeted Regeneration

2022

Glossary

ACLF – Acute-on-chronic liver failure–

ACTA2 – actin protein

ADA – Anti-drug antibodies

AH – Alcoholic hepatitis

ALT – Alanine Aminotransferase

AMD – Age-related macular degeneration

ASGR1 – Asiaglycoprotein receptor 1

AST – Aspartate aminotransferase

AT1/AT2 – Alveolar type epithelial cells

BW – Body weight

COPD – Chronic Obstructive Pulmonary Disease

DC – Dendritic cell

DME – Diabetic macular edema

DSS – Dextran sodium sulfate

EtOH – Ethyl alcohol

FSGS – Focal segmental glomerulosclerosis

Fzd – Frizzled

GFP – Green fluorescence protein

GI – Gastrointestinal

GLP – Good laboratory practice

HNF alpha - Hepatocyte nuclear factor 4 alpha

IBD – inflammatory Bowel Disease

IgG – Immunoglobulin G

IPF – Idiopathic pulmonary fibrosis

IND – Investigational new Drug

INR – International normalized ratio

IV – Intravenous

KO – Knock-out model

LG – Lacrimal gland

Lille – Modeling tool for predicting mortality in patients with alcoholic hepatitis who are not responding to steroid therapy

Lrp Lipoprotein receptor-related protein

MELD – Model for end-stage liver disease score

MOA – Mechanism of action

PD – Pharmacodynamics

Pg – Picogram

Mg – Milligrams

MS – Multiple sclerosis

PIPE – Private investment in public equity

PK – Pharmacokinetic

SAD – Single ascending dose

SC – Subcutaneous

MAD – Multiple ascending dose

RPE – Retinal pigment epithelium

SAH – severe alcoholic hepatitis

SOC – Standard of care

SWAP – Surrozen Wnt signal activating proteins

SWEETS – Surrozen Wnt enhancer engineered for tissue specificity

TAA – Thioacetamide

UC – Ulcerative colitis; Mod-Sev UC – Moderate to Severe UC

UC-100 – A score of a composite disease activity index for drug development in UC

VHH – Single variable domain on a heavy chain (VHH) antibodies

