



A New Path to Medicine

Ladenburg Thalmann 2019 Healthcare Conference

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September 24, 2019

Forward-Looking Statements

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aTyr Pharma

Company Overview

Accelerating Value Creation from New Biology

Platform of New Biology:

Discovery pipeline of novel therapeutic candidates based on proprietary knowledge of extracellular functions of tRNA synthetases (~300 protein compositions patented)

Lead Product Candidate: ATYR1923

Engineered, long acting, protein therapeutic, derived from the HARS gene, for the treatment of pulmonary sarcoidosis and other interstitial lung diseases

\$2-3b⁽¹⁾ global opportunity

Financials:

Cash, cash equivalents and investments at \$42.4m as of 6/30/2019

April 2019: \$5m raise with Federated and Dr. Paul Schimmel, board member, at market, no discount or warrants

Clinical Milestones:

Initiated P1b/2a Trial – Q4 2018

- ☐ Interim Safety – Q4 2019
- ☐ Final Results – mid-2020⁽²⁾

(1) aTyr estimates for inflammatory ILD: Pulmonary Sarcoidosis, CHP, CTD-ILD; excludes IPF

(2) Dependent on patient enrollment

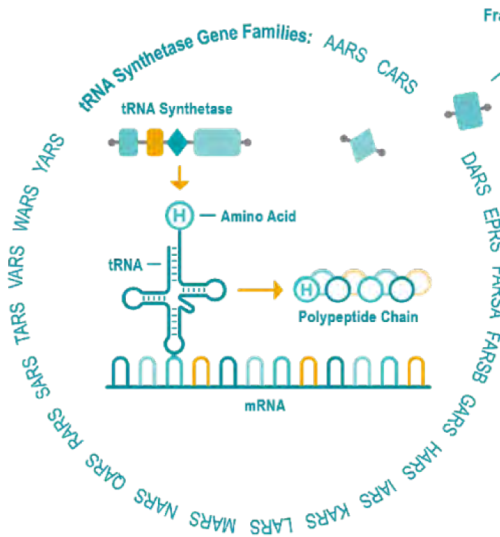
Development Pipeline

PROGRAM	DISEASES	DISCOVERY	PRECLINICAL	PHASE 1	PHASE 1B/2	PHASE 2/3
ATYR1923	Pulmonary Sarcoidosis					
	Chronic Hypersensitivity Pneumonitis (CHP)					
	Connective Tissue Disease ILD (CTD-ILD)					
tRNA Synthetase Candidates	Undisclosed		CSL Behring			
NRP2 Candidates	Undisclosed					

Extracellular tRNA Synthetase Biology

INTRACELLULAR

Catalyze Protein Synthesis



EXTRACELLULAR

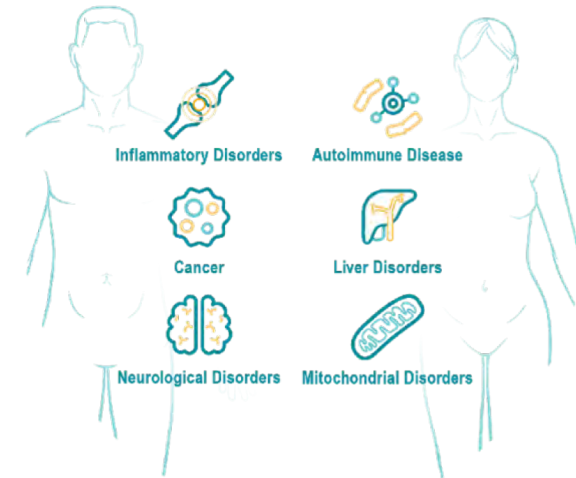
Secreted in Circulation and Tissue to Regulate Diverse Pathways

Fragments and Splice Variants



PHYSIOLOGICAL

Pathway Disruption Associated with Disease



CSL Behring Collaboration

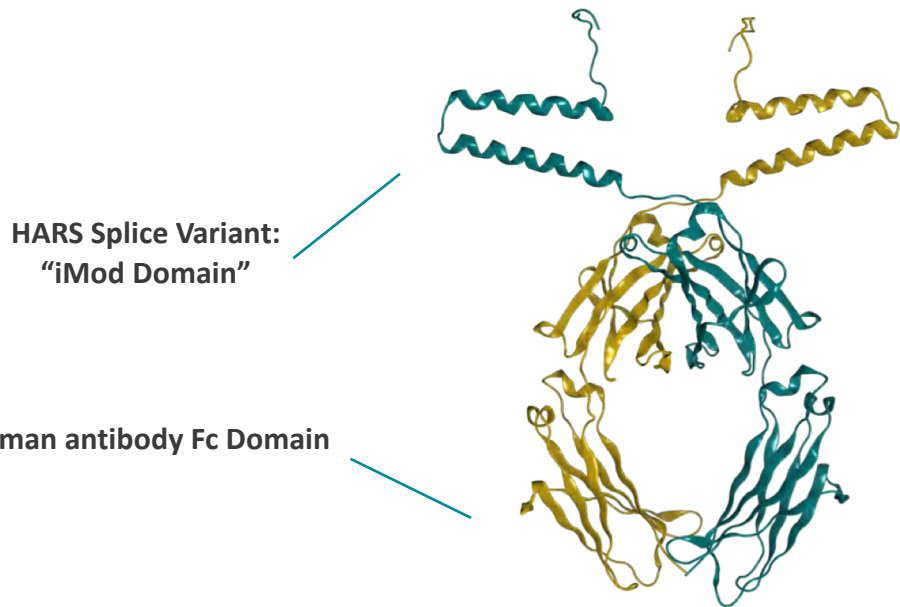
Goal	<ul style="list-style-type: none">Identify new IND candidates from up to four tRNA synthetases from aTyr's proprietary pipeline of novel proteins (non-HARS derived)
Terms	<ul style="list-style-type: none">CSL Behring to fund all R&D costsaTyr eligible for up to \$17m in option fees if CSL Behring advances all four programs (\$4.25m per synthetase program)aTyr grants CSL Behring an option to negotiate licenses for worldwide rights to each IND candidate that emerges from the collaboration
About CSL	<ul style="list-style-type: none">CSL Behring is a global biotherapeutics leader specializing in immunology, hematology and other rare and serious medical conditionsCSL Behring employs >22,000 people globally, and delivers its therapies to more than 60 countries
Status	<ul style="list-style-type: none">aTyr received first phase of funding totaling \$630k, and of that recognized \$94k of collaboration revenue in Q2 2019



ATYR1923

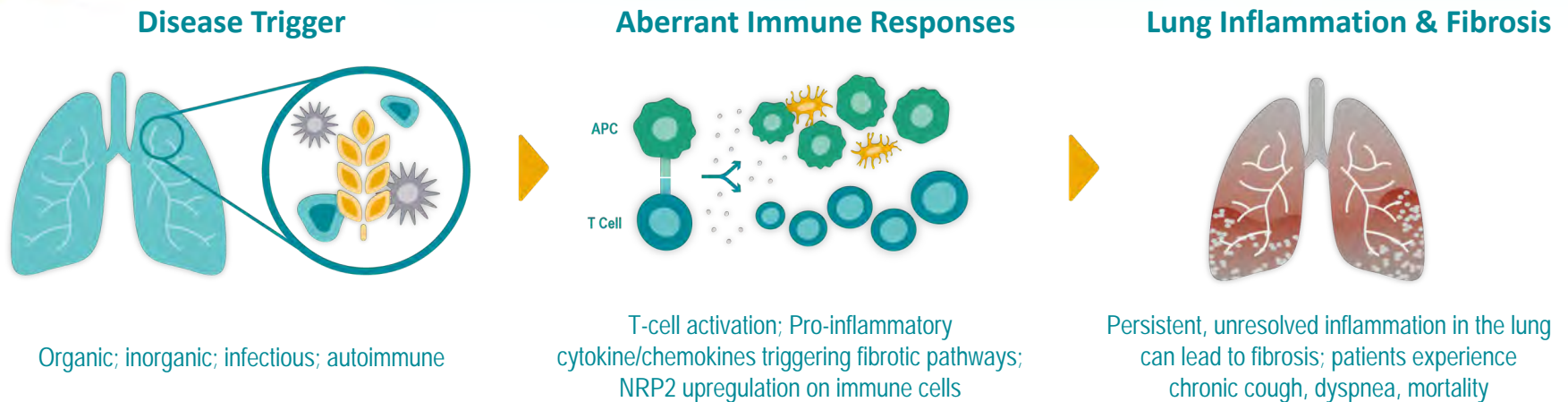
For the Treatment of
Pulmonary Sarcoidosis

ATYR1923: Novel Engineered Protein Therapeutic

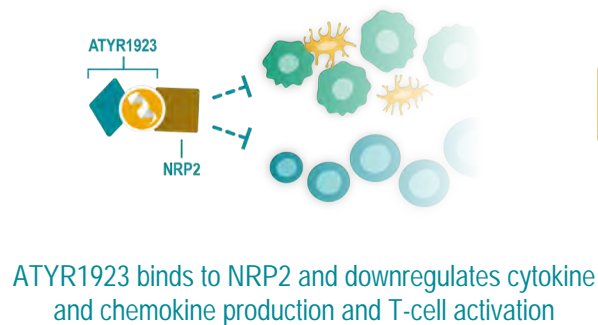


- iMod Domain of HARS enriched in the human lung
- Inhibits human T cell activation/cytokine release
- Binds selectively to Neuropilin-2 (NRP2)
- Regulates a number of immune cell-types, including: T cells, Neutrophils, Macrophages, Dendritic cells
- iMod Domain fused to Fc Domain to extend half-life
- Once-monthly IV dosing regimen

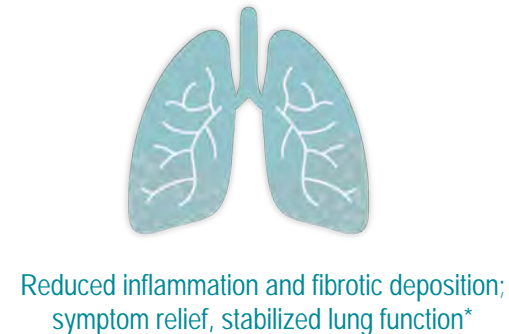
ATYR1923 Mechanism of Action in ILD



ATYR1923 Dampens Immune Responses



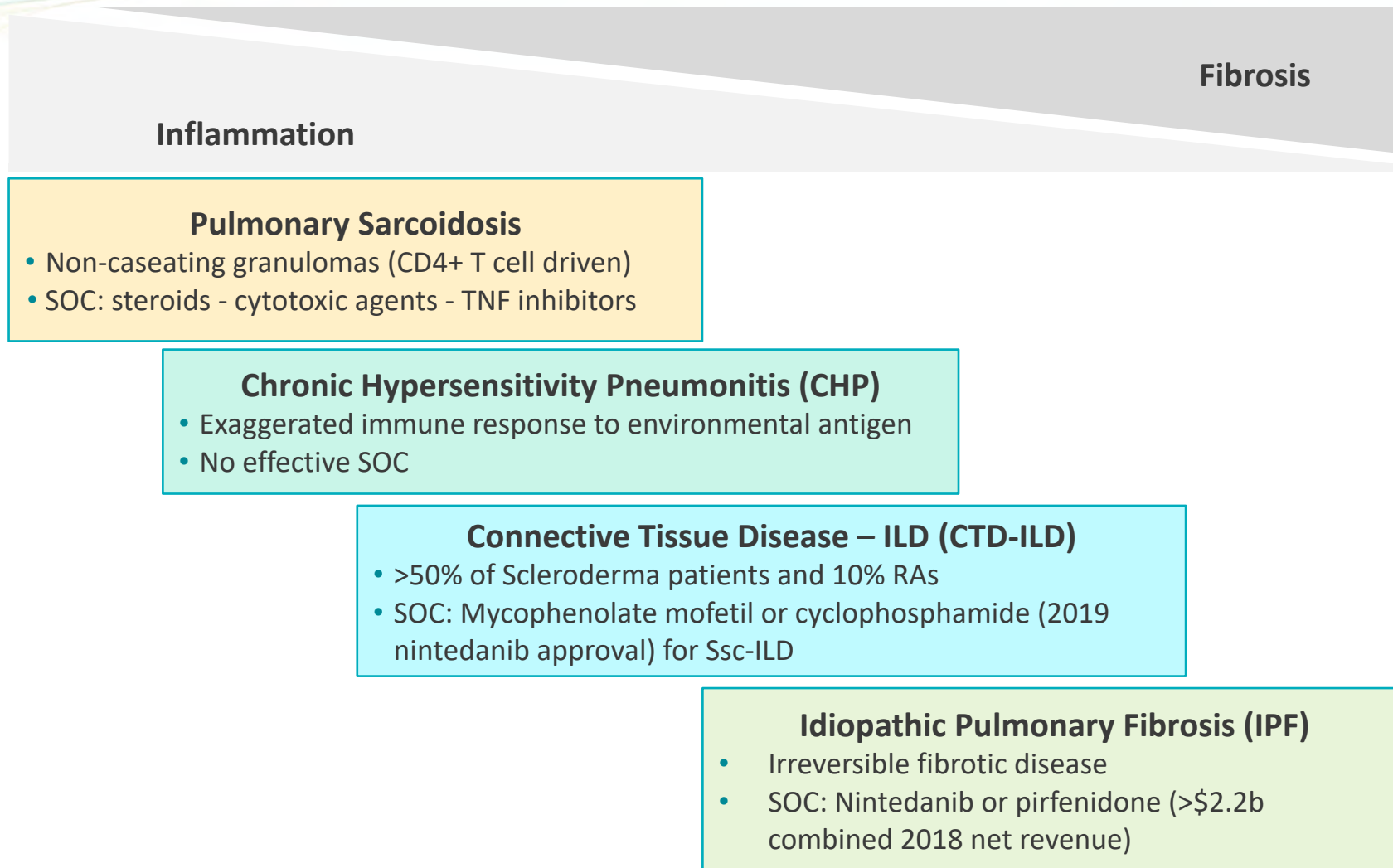
Stabilized Lung



Pre-Clinical Translational Data Supports ILD Development

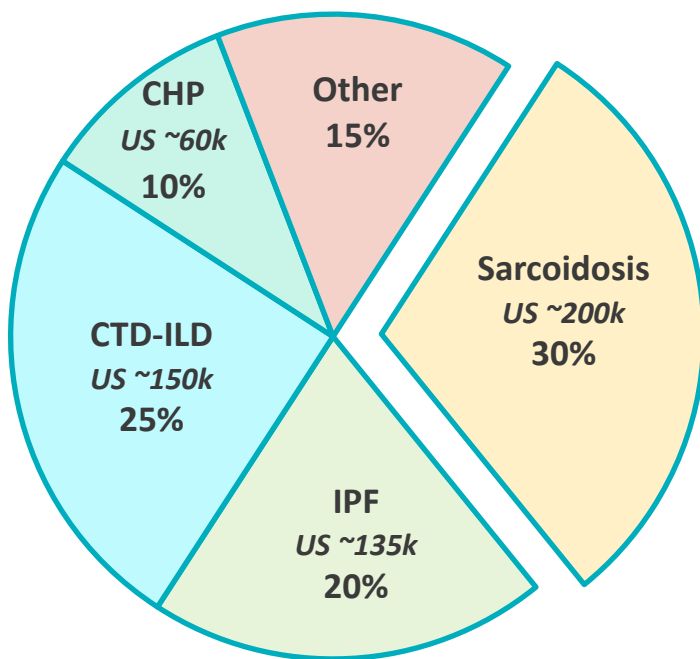
Bleomycin-Induced Lung Injury (IPF) – Mouse	<ul style="list-style-type: none">• ATYR1923 reduced fibrosis and inflammation• Comparator: pirfenidone• Presented at ATS, May 2017
Bleomycin-Induced Lung Injury (IPF) – Rat	<ul style="list-style-type: none">• ATYR1923 returned lung function to normal and reduced fibrosis and inflammation• Comparator: nintedanib• Presented at ATS, May 2018
Sclerodermatous chronic-graft vs host disease (SSc-ILD) – Mouse	<ul style="list-style-type: none">• ATYR1923 reduced lung and skin fibrosis• Comparator: nintedanib• Presented at Scleroderma Foundation Patient Conference, July 2018
SSc-cGVHD (SSc-ILD); <i>P. acnes</i> (Sarcoidosis); <i>S. rectivirgula</i> (CHP); SKG (Ra-ILD) – Mouse	<ul style="list-style-type: none">• ATYR1923 demonstrated stage-dependent anti-inflammatory and anti-fibrotic effect in various experimental models of ILD• Comparator: various• Presented at ATS, May 2019

ILDs Share Persistent Immune Engagement

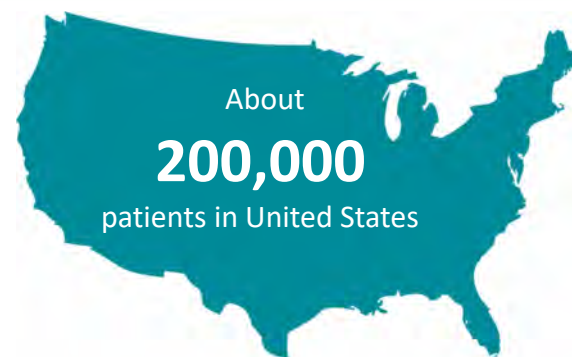


Sarcoidosis: A Major Form of ILD

ILD Patient Distribution



\$2-3b Global Opportunity⁽¹⁾



50% require systemic therapy



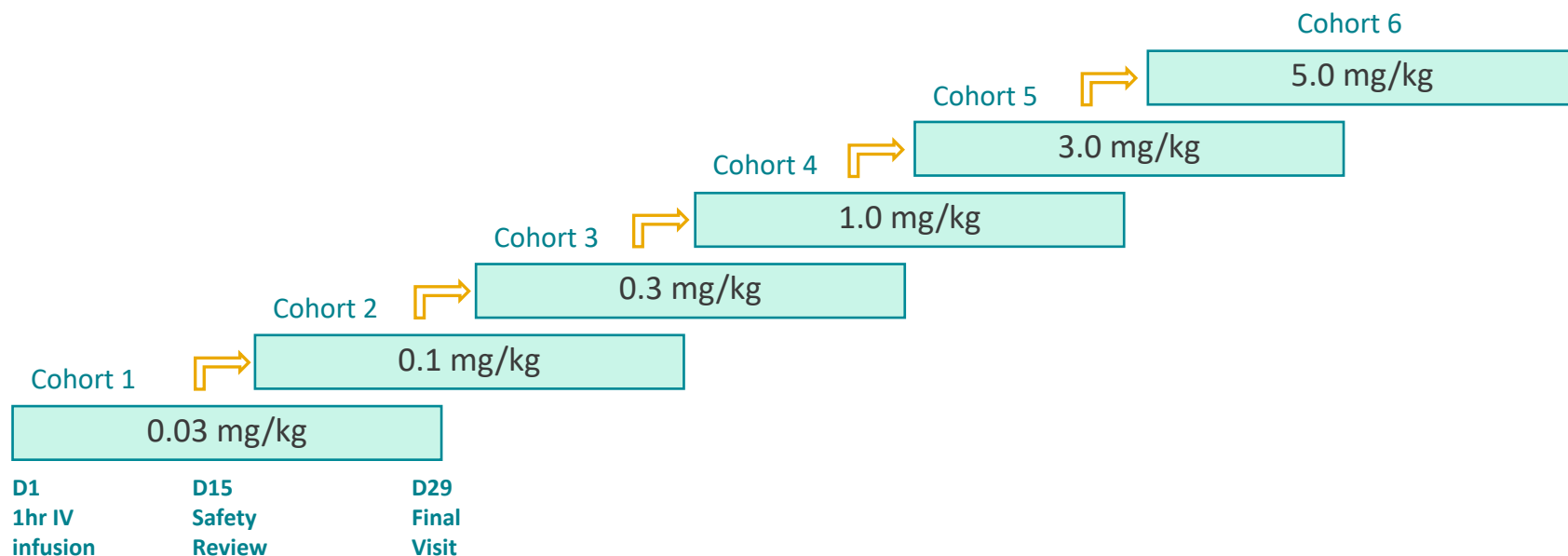
30% with chronic progressive disease despite currently available treatment



PK Profile Supports Potential Once-Monthly Dosing

Phase 1 Healthy Volunteer Study Completed in Australia

- Positive data announced in June 2018
- Randomized, double-blind, placebo-controlled, single ascending dose (N=36 HVs)
- ATYR1923 was generally well-tolerated with no significant adverse events



ATYR1923 Phase 1b/2a Study in Pulmonary Sarcoidosis

Design	<ul style="list-style-type: none">• Randomized, double-blind, placebo-controlled, multiple ascending dose
Population	<ul style="list-style-type: none">• Histologically confirmed pulmonary sarcoidosis• Requiring ≥ 10 mg prednisone (steroid) treatment; capable of steroid taper• Symptomatic/active disease at baseline by ^{18}F-FDG-PET/CT, Pulmonary Function Tests
Dosing	<ul style="list-style-type: none">• 3 sequential cohorts, 12 patients each• 2:1 randomization• ATYR1923 doses: 1.0, 3.0, and 5.0 mg/kg
Duration	<ul style="list-style-type: none">• 24-week study period• Steroid taper phase down to 5.0 mg by week 8• 16-week maintenance phase
Sites	<ul style="list-style-type: none">• Up to ~15 leading pulmonary sarcoidosis centers• Collaboration with the Foundation for Sarcoidosis Research

ATYR1923 Phase 1b/2a Study Endpoints

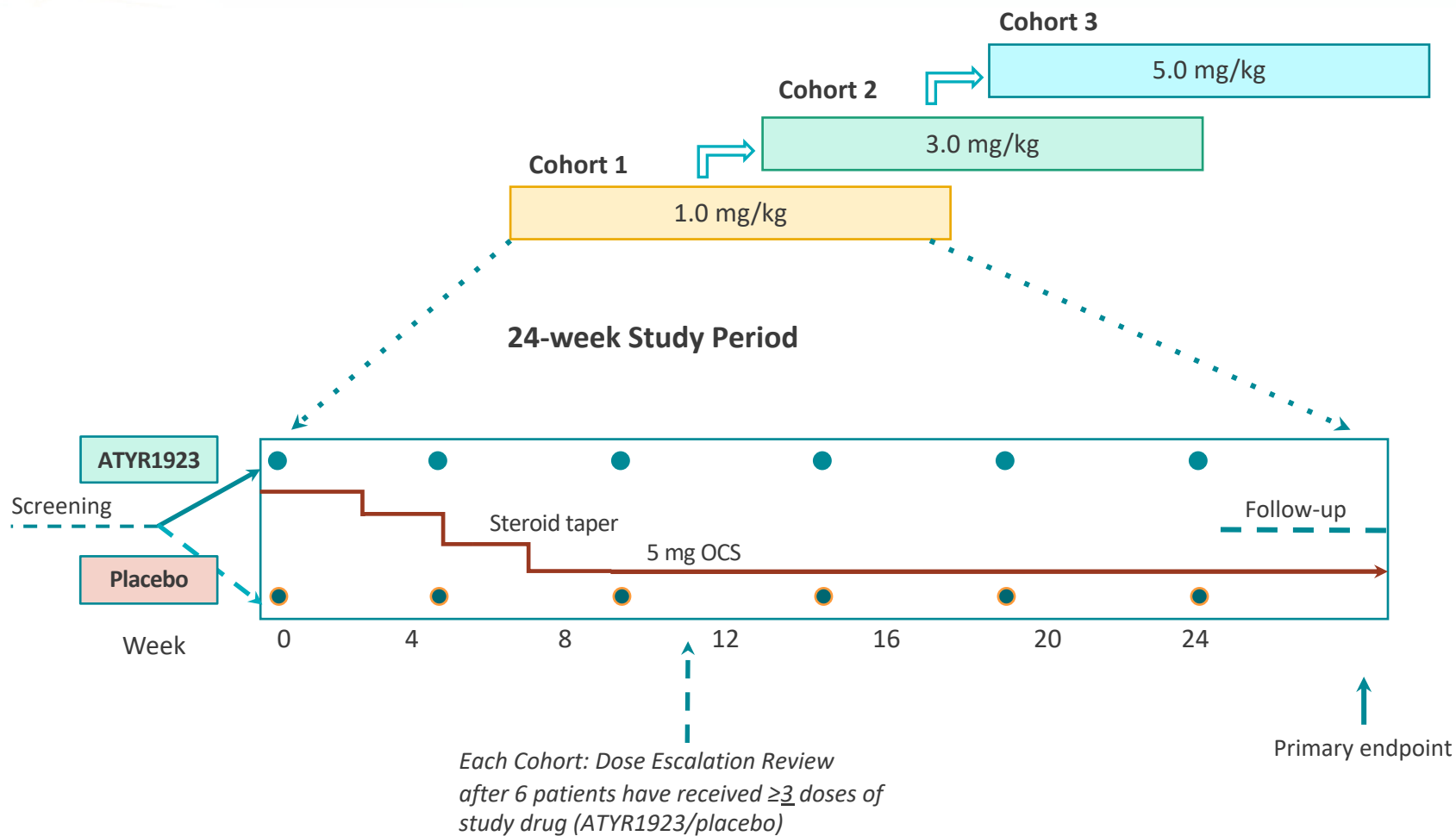
Primary

- Safety and tolerability of multiple ascending IV ATYR1923 doses

Secondary

- Steroid-sparing effect
- Immunogenicity
- Pharmacokinetics (PK)
- Exploratory efficacy measures: FDG-PET/CT imaging; Lung function (FVC); Serum biomarkers; Health-related quality of life scales

Phase 1b/2a Study Schema



ATYR1923 Phase 1b/2a Study in Pulmonary Sarcoidosis

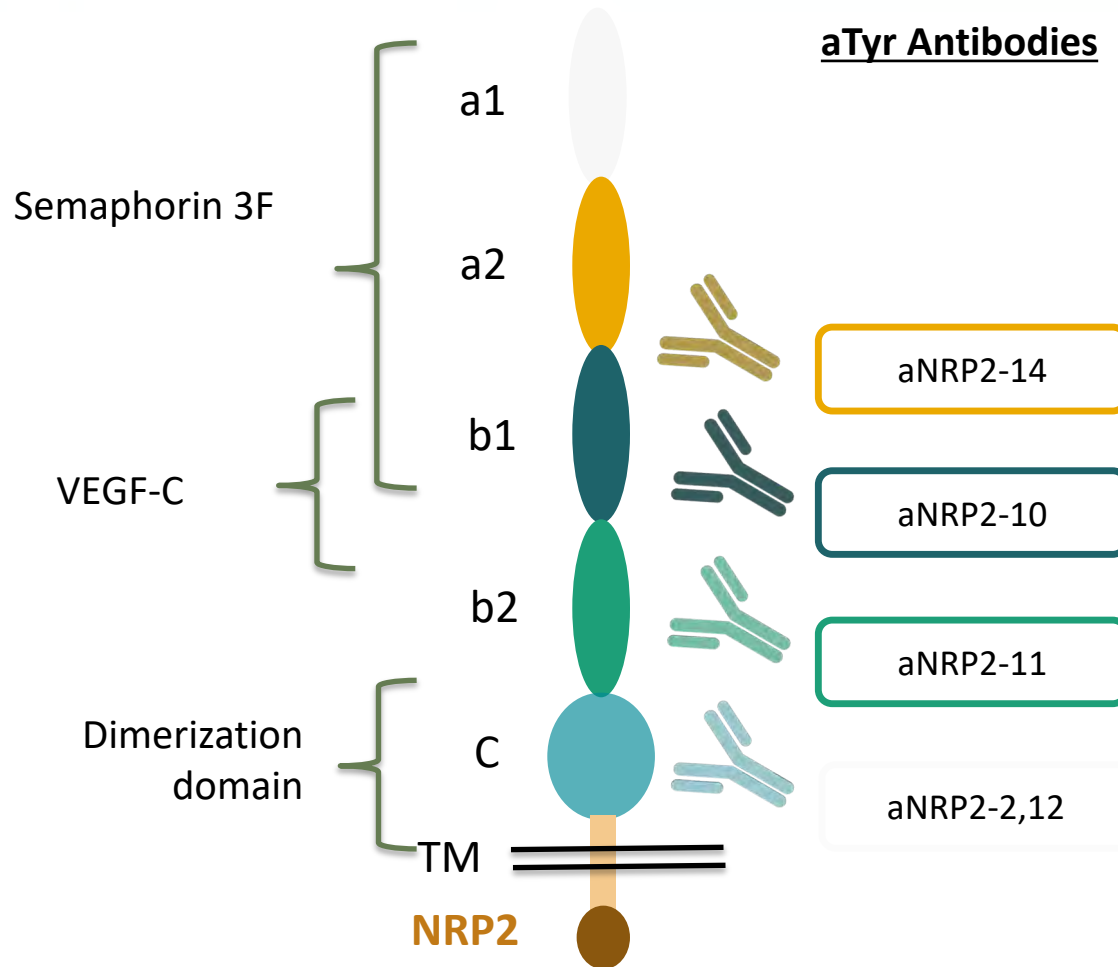
Status	<ul style="list-style-type: none">• Patient enrollment ongoing• Evaluating additional sites
Timelines	<ul style="list-style-type: none">• Interim safety data: Q4 2019• Study completion: mid-2020⁽¹⁾
Possible Future Development	<ul style="list-style-type: none">• Registrational trial in Pulmonary Sarcoidosis• Initiate P2 studies in other types of interstitial lung disease (e.g. CTD-ILD; CHP)

(1) Dependent on patient enrollment

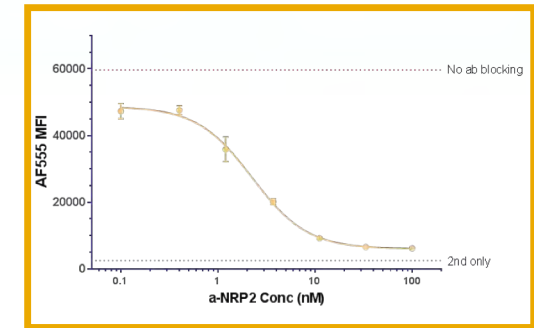


NRP2 Biology

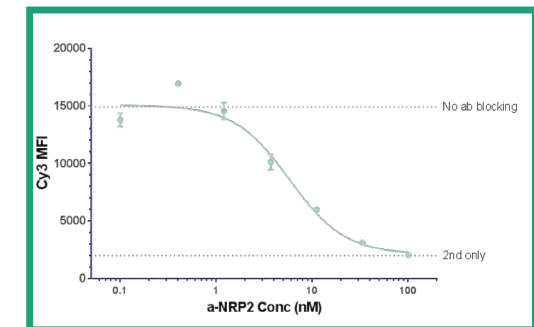
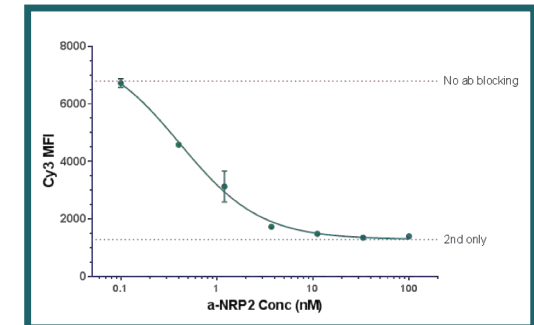
aTyr NRP2 Blocking Antibodies



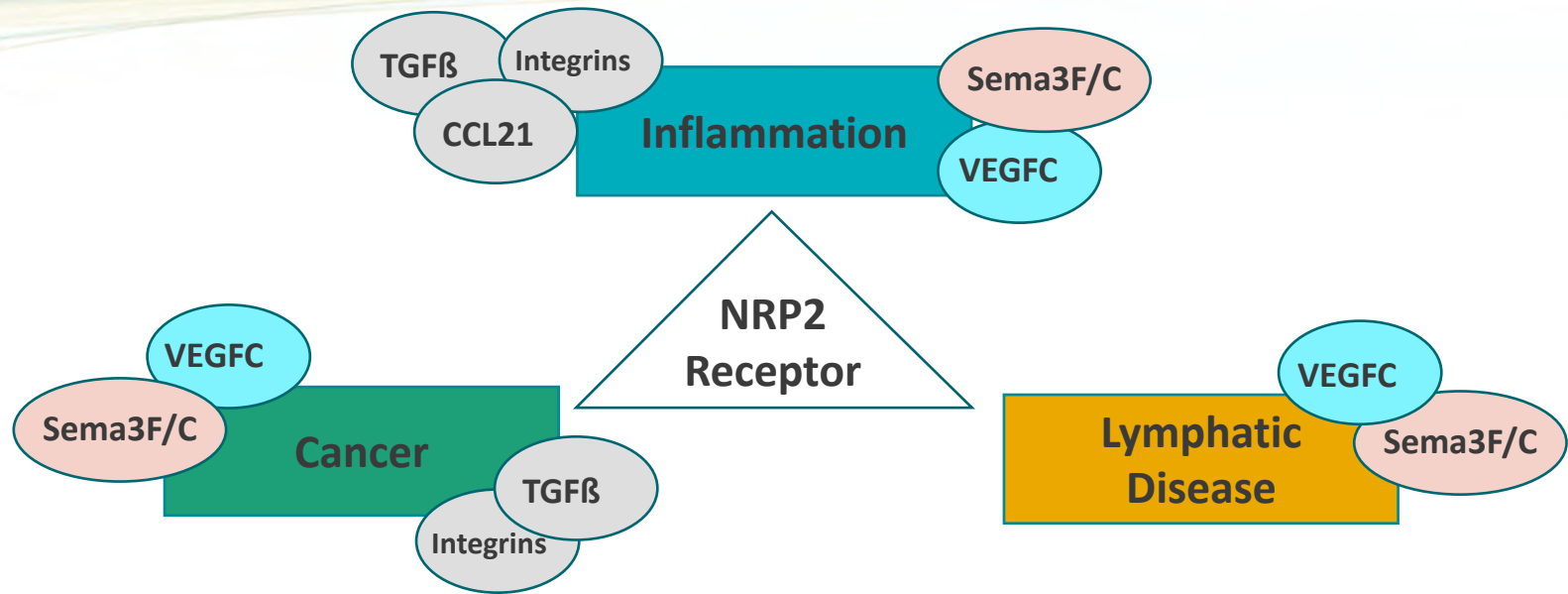
Sema3F Blocking



VEGF-C Blocking



NRP2 Receptor Biology Associated with Diverse Pathways



- Implicated in cancer, inflammation and lymphatic disease
- Co-receptors for semaphorins and VEGF family molecules
- Overexpressed in various tumors, tumor expression linked to poor prognosis
- Critical for cancer cell migration, metastasis, EMT, lymphangiogenesis



aTyr Pharma

Company Value Drivers

Upcoming Catalysts

ATYR1923

- ❑ Interim Phase 1b/2a safety data Q4 2019
- ❑ Phase 1b/2a results mid-2020⁽¹⁾
- ❑ Potential expansion into Phase 2 studies for CHP and CTD-ILD

CSL R&D

- ❑ aTyr eligible for up to \$17m in option fees
- ❑ Option granted to CSL to negotiate licenses for worldwide rights to each IND candidate that emerges from the collaboration

NRP2 Antibody Candidates

- ❑ Potential new pipeline opportunities through academic and industry collaborations

(1) Dependent on patient enrollment

Building Value...for Patients and Shareholders

- ✓ Platform of new biology
 - ✓ tRNA synthetase biology
 - ✓ ~300 protein compositions patented
 - ✓ NRP2 antibody program
- ✓ Robust clinical program: ATYR1923
 - ✓ Understanding of MOA
 - ✓ Translational studies in multiple ILD models
 - ✓ Phase 1b/2a clinical study in pulmonary sarcoidosis
- ✓ Supported by top tier investors
- ✓ Cash, cash equivalents, and investment at \$42.4m as of 6/30/2019



Thank You