



A New Path to Medicine

Ladenburg Thalmann 2019 Healthcare Conference
Sanjay S. Shukla, M.D., M.S., President & CEO

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⁾

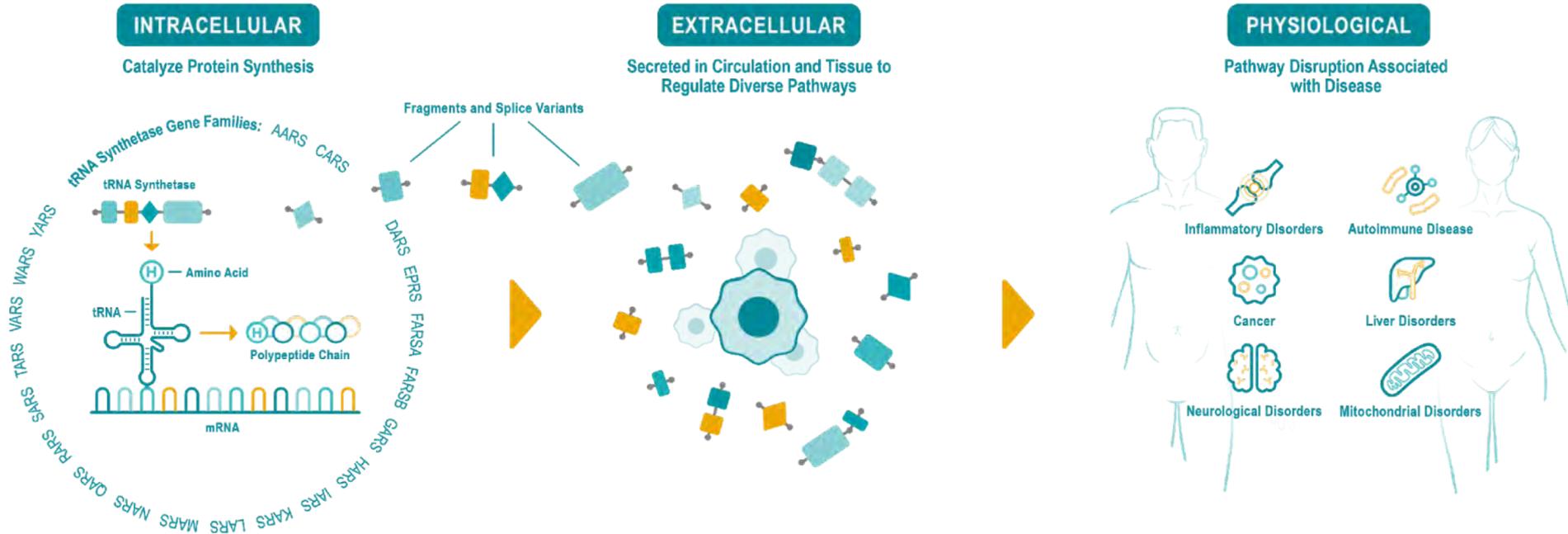
(1) aTyr estimates for inflammatory ILD: Pulmonary Sarcoidosis, CHP, CTD-ILD; excludes IPF

(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					
NRP2 Candidates	Undisclosed					

Extracellular tRNA Synthetase Biology



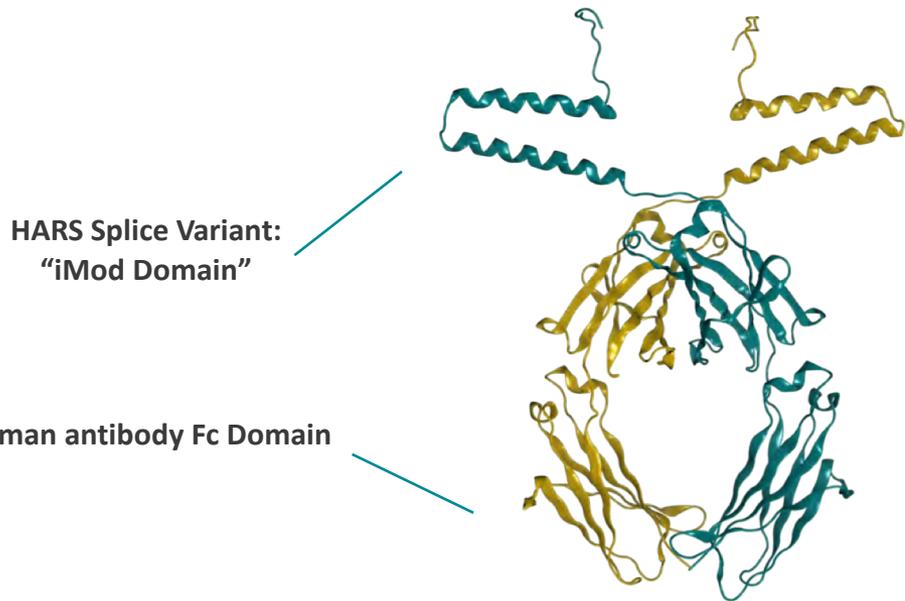
CSL Behring Collaboration

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 costs• aTyr 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 conditions• CSL 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

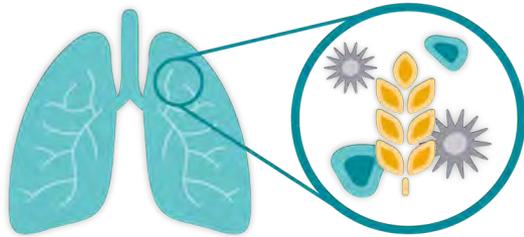
ATYR1923: Novel Engineered Protein Therapeutic



- 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

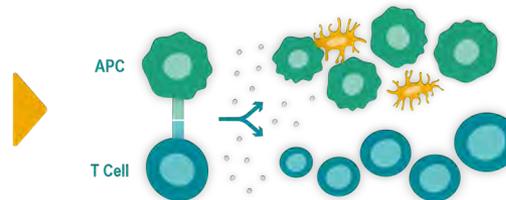
ATYR1923 Mechanism of Action in ILD

Disease Trigger



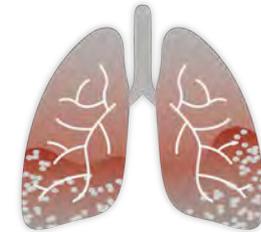
Organic; inorganic; infectious; autoimmune

Aberrant Immune Responses



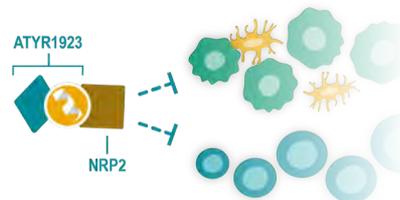
T-cell activation; Pro-inflammatory cytokine/chemokines triggering fibrotic pathways; NRP2 upregulation on immune cells

Lung Inflammation & Fibrosis



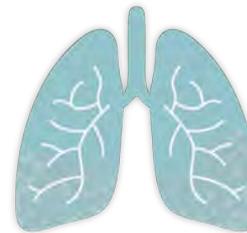
Persistent, unresolved inflammation in the lung can lead to fibrosis; patients experience chronic cough, dyspnea, mortality

ATYR1923 Dampens Immune Responses



ATYR1923 binds to NRP2 and downregulates cytokine and chemokine production and T-cell activation

Stabilized Lung



Reduced inflammation and fibrotic deposition; symptom relief, stabilized lung function*

Pre-Clinical Translational Data Supports ILD Development

Bleomycin-Induced Lung Injury (IPF) – Mouse

- ATYR1923 reduced fibrosis and inflammation
- Comparator: pirfenidone
- Presented at ATS, May 2017

Bleomycin-Induced Lung Injury (IPF) – Rat

- ATYR1923 returned lung function to normal and reduced fibrosis and inflammation
- Comparator: nintedanib
- Presented at ATS, May 2018

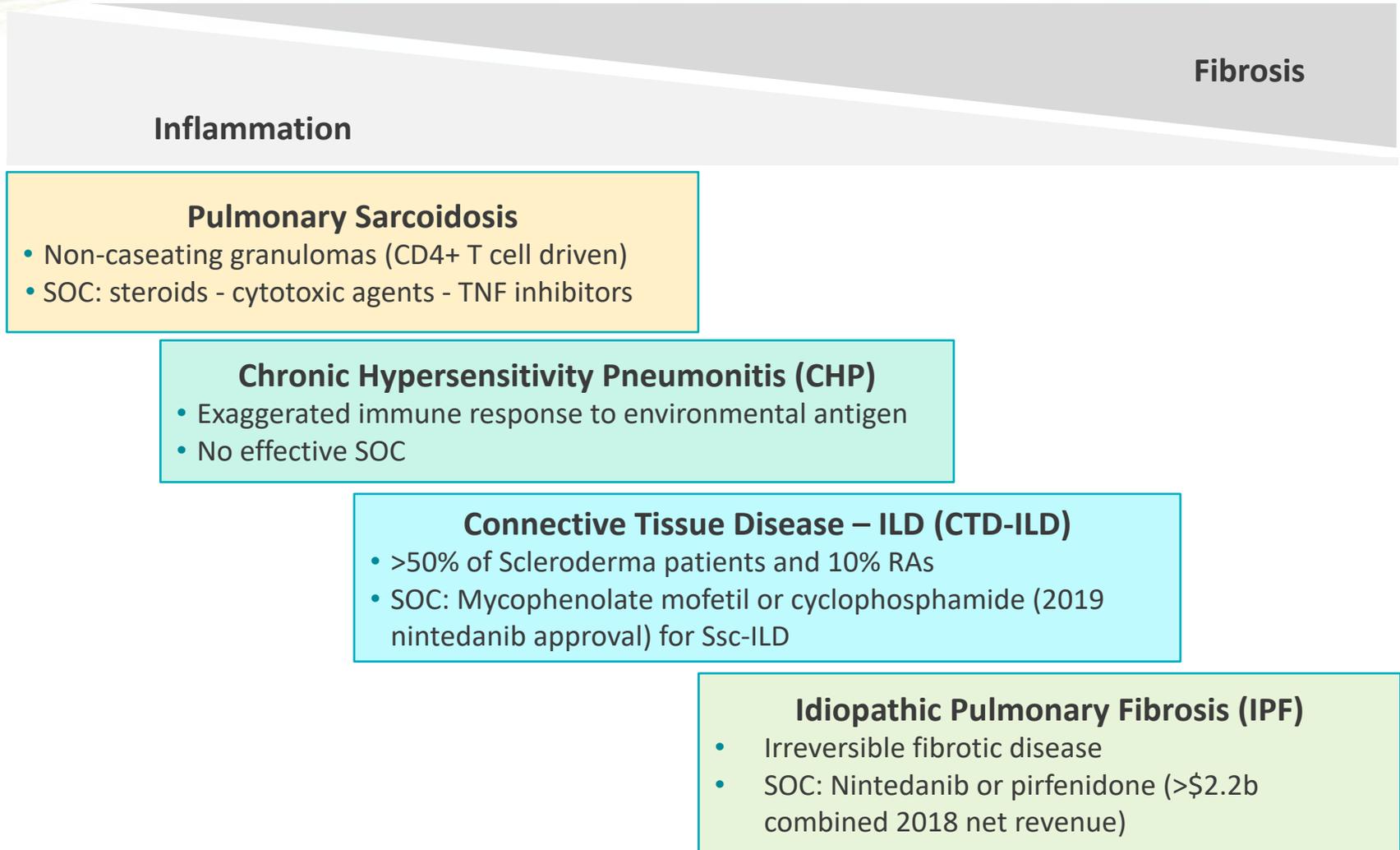
Sclerodermatous chronic-graft vs host disease (SSc-ILD) – Mouse

- ATYR1923 reduced lung and skin fibrosis
- Comparator: nintedanib
- Presented at Scleroderma Foundation Patient Conference, July 2018

SSc-cGVHD (SSc-ILD); *P. acnes* (Sarcoidosis); *S. rectivirgula* (CHP); SKG (Ra-ILD) – Mouse

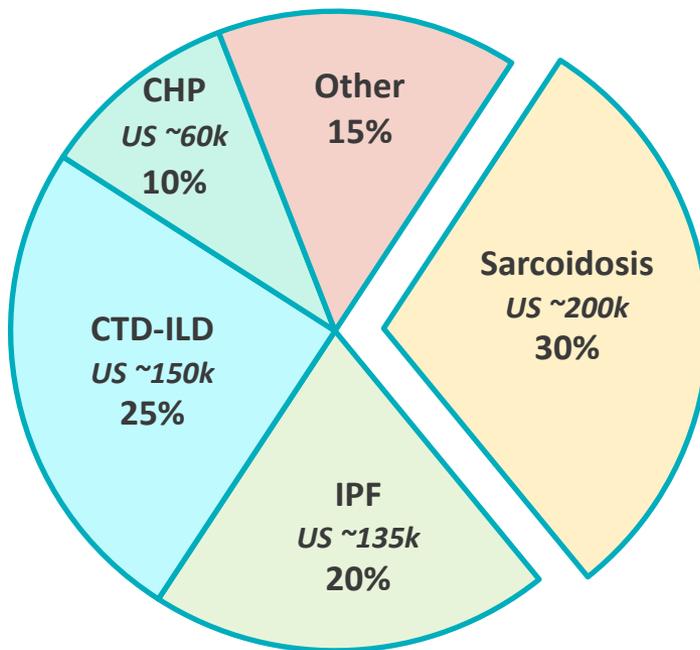
- ATYR1923 demonstrated stage-dependent anti-inflammatory and anti-fibrotic effect in various experimental models of ILD
- Comparator: various
- Presented at ATS, May 2019

ILDs Share Persistent Immune Engagement



Sarcoidosis: A Major Form of ILD

ILD Patient Distribution



\$2-3b Global Opportunity⁽¹⁾



50% require systemic therapy



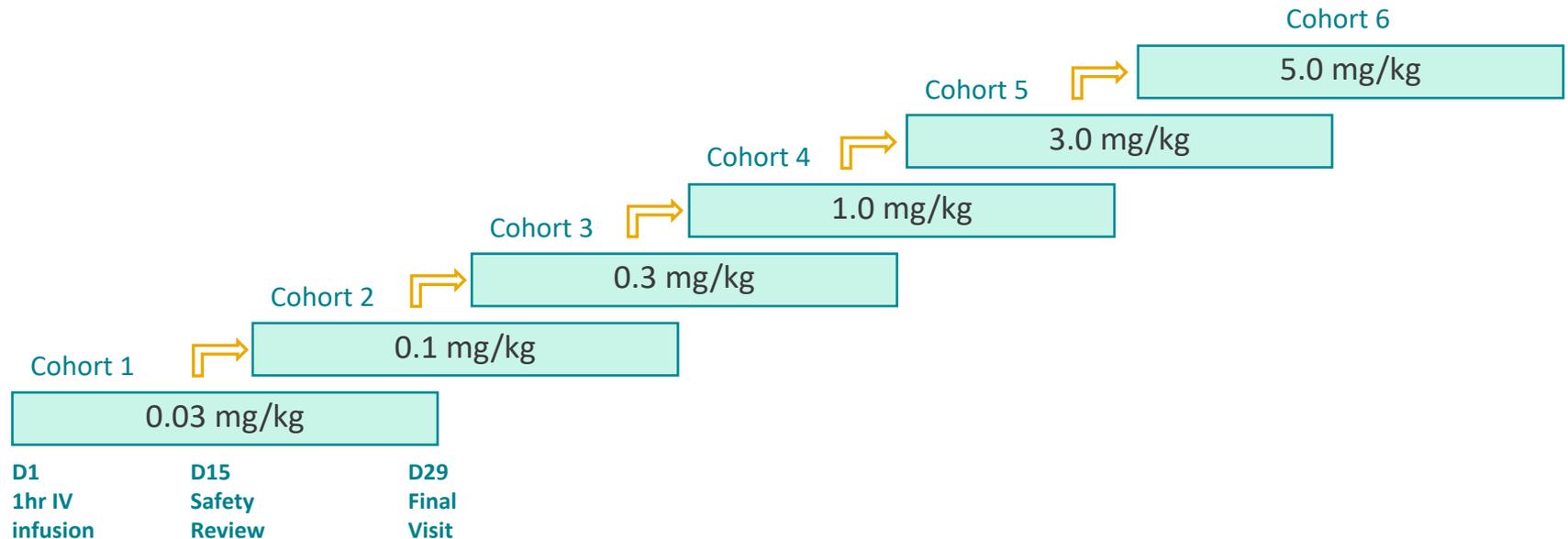
30% with chronic progressive disease despite currently available treatment



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

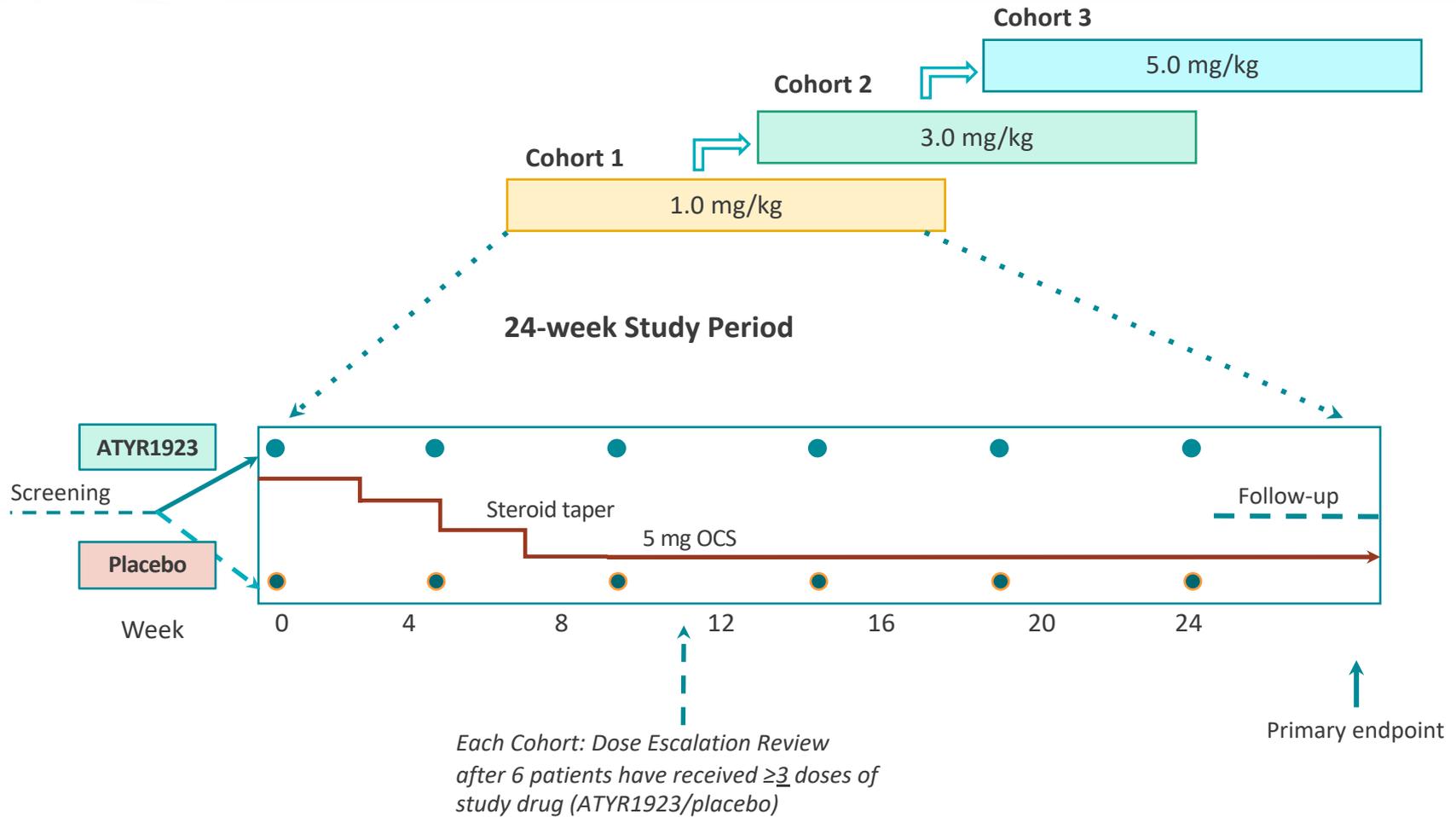
Primary

- 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

- Patient enrollment ongoing
- Evaluating additional sites

Timelines

- Interim safety data: Q4 2019
- Study completion: mid-2020⁽¹⁾

Possible Future Development

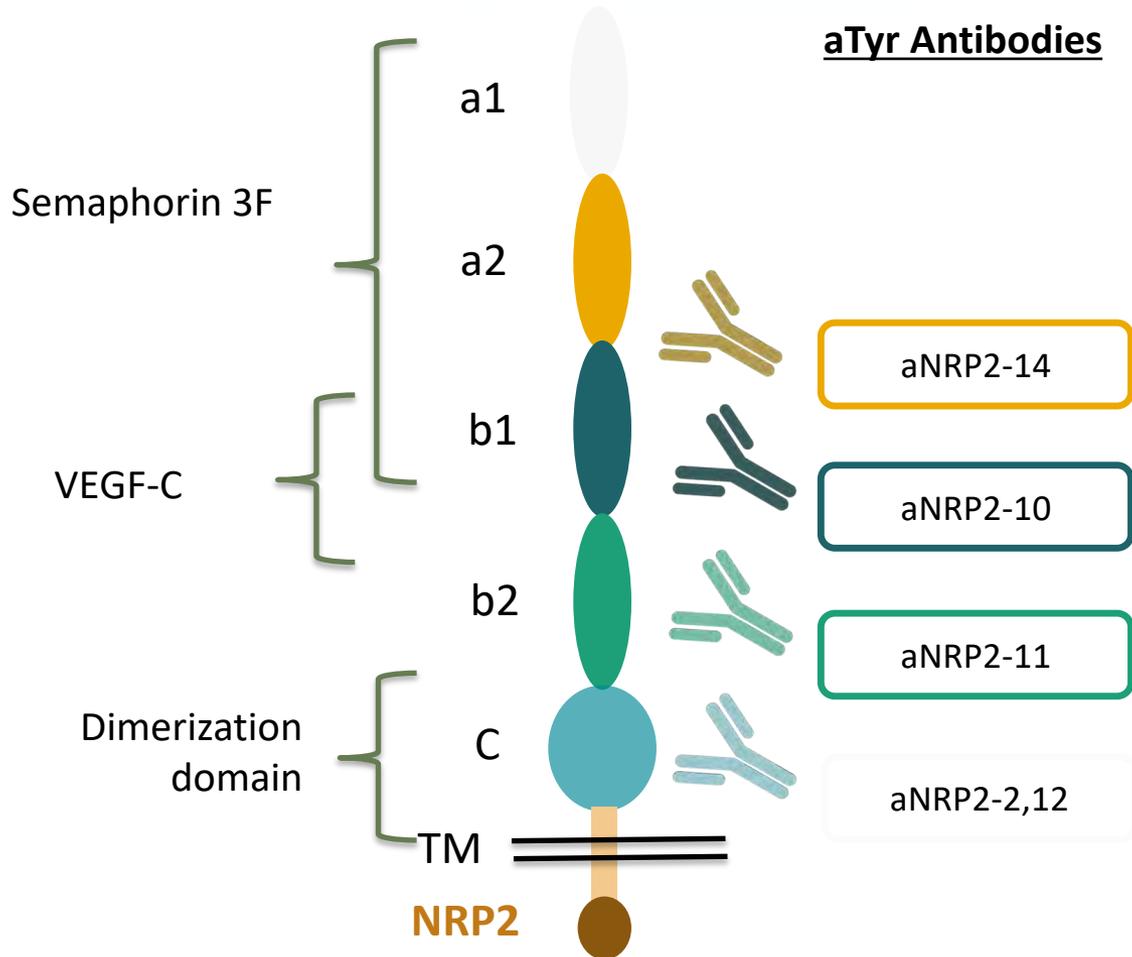
- Registrational trial in Pulmonary Sarcoidosis
- Initiate P2 studies in other types of interstitial lung disease (e.g. CTD-ILD; CHP)

(1) Dependent on patient enrollment

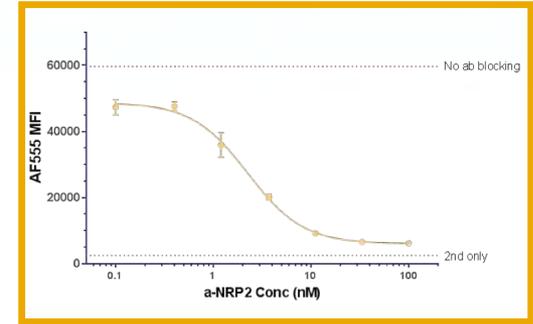


NRP2 Biology

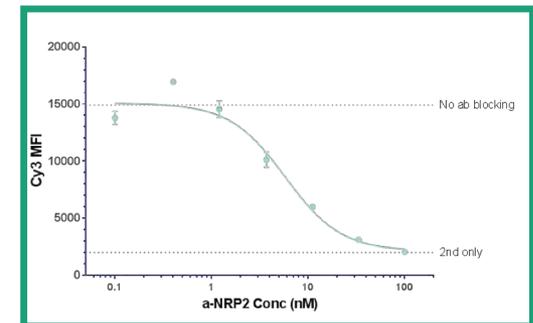
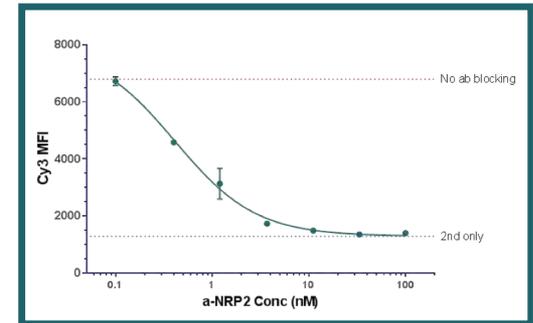
aTyr NRP2 Blocking Antibodies



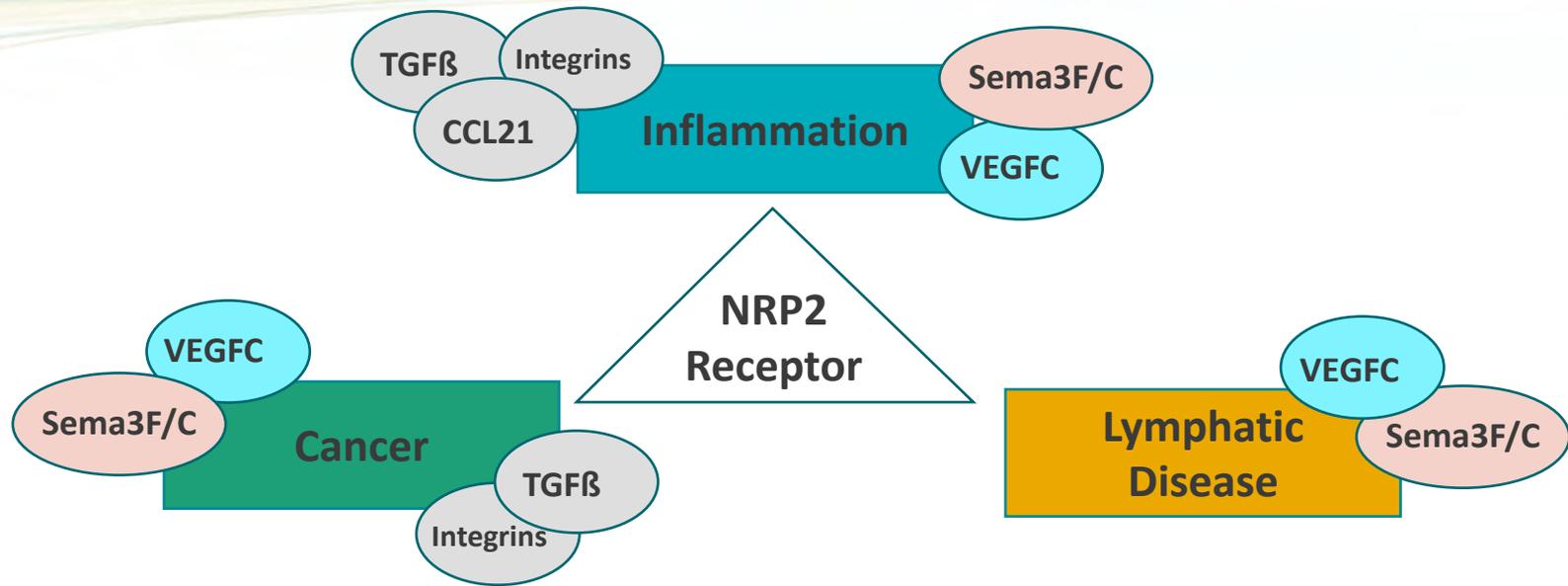
Sema3F Blocking



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