

**UNITED STATES  
SECURITIES AND EXCHANGE COMMISSION**  
Washington, D.C. 20549

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**FORM 8-K**

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**CURRENT REPORT**

**Pursuant to Section 13 or 15(d) of the  
Securities Exchange Act of 1934**

**Date of Report (Date of earliest event reported): January 11, 2016**

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**ATYR PHARMA, INC.**

(Exact name of registrant as specified in its charter)

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**Delaware**  
(State or other jurisdiction of  
incorporation)

**001-37378**  
(Commission  
File Number)

**20-3435077**  
(I.R.S. Employer  
Identification No.)

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**3545 John Hopkins Court, Suite #250  
San Diego, CA 92121**

(Address of principal executive offices, including zip code)

**(858) 731-8389**

(Registrant's telephone number, including area code)

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Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

- Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)
  - Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)
  - Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))
  - Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))
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**Item 7.01 Regulation FD Disclosure.**

aTyr Pharma, Inc. (the “Company”) intends to use an investor presentation to conduct meetings with investors, stockholders and analysts and at investor conferences, and which the Company intends to place on its website. A copy of the presentation materials is attached hereto as Exhibit 99.1. The Company does not undertake to update the presentation materials.

The information under this Item 7.01, including Exhibit 99.1, is being furnished and shall not be deemed “filed” for the purposes of Section 18 of the Securities and Exchange Act of 1934, as amended, or the Exchange Act, or otherwise subject to the liabilities of that section, nor shall they be deemed incorporated by reference into any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as expressly set forth by specific reference in such filing

**Item 9.01 Financial Statements and Exhibits.**

(d) Exhibits

<b>Exhibit No.</b>	<b>Description</b>
99.1	Corporate Presentation Materials of aTyr Pharma, Inc. dated January 2016

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## SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

Date: January 11, 2016

**aTyr Pharma, Inc.**

By: /S/ JOHN D. MENDLEIN

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John D. Mendlein, Ph.D.

Chief Executive Officer

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EXHIBIT INDEX

**Exhibit No.**

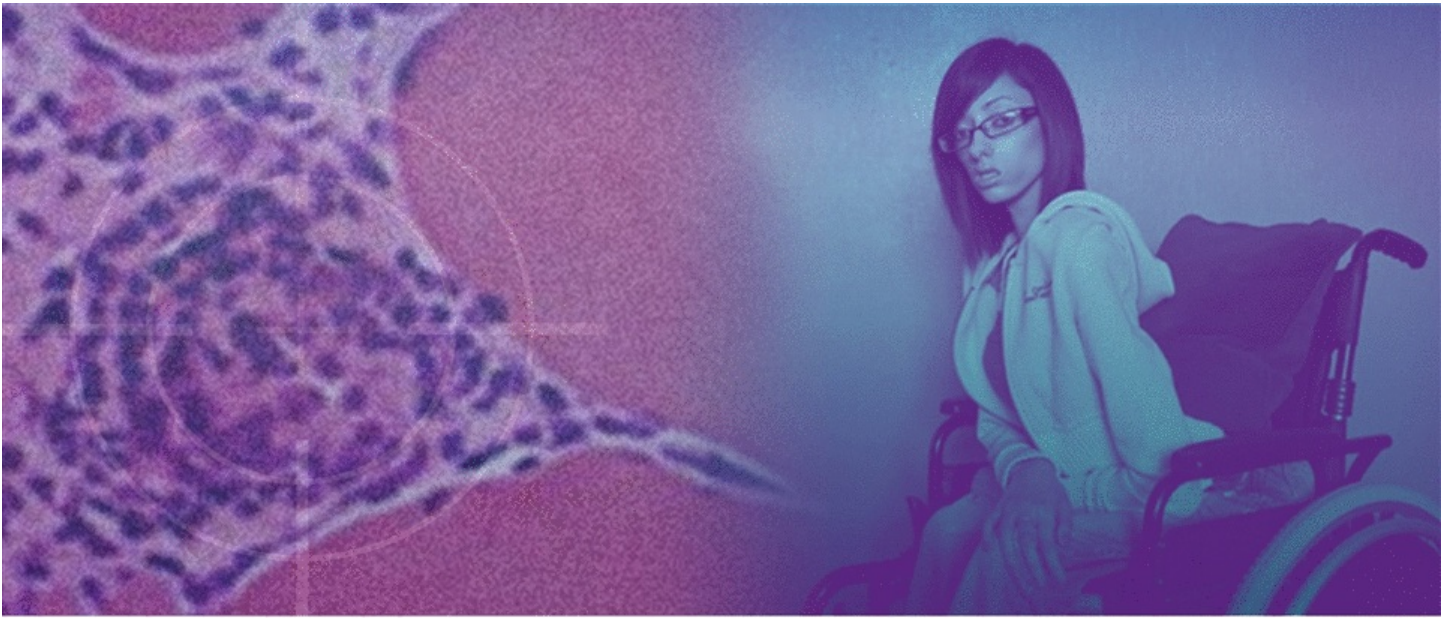
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**Description**

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99.1

Corporate Presentation Materials of aTyr Pharma, Inc. dated January 2016



*New Promise for Rare Disease Patients  
with Immune & Fibrotic Pathologies*

January 2016



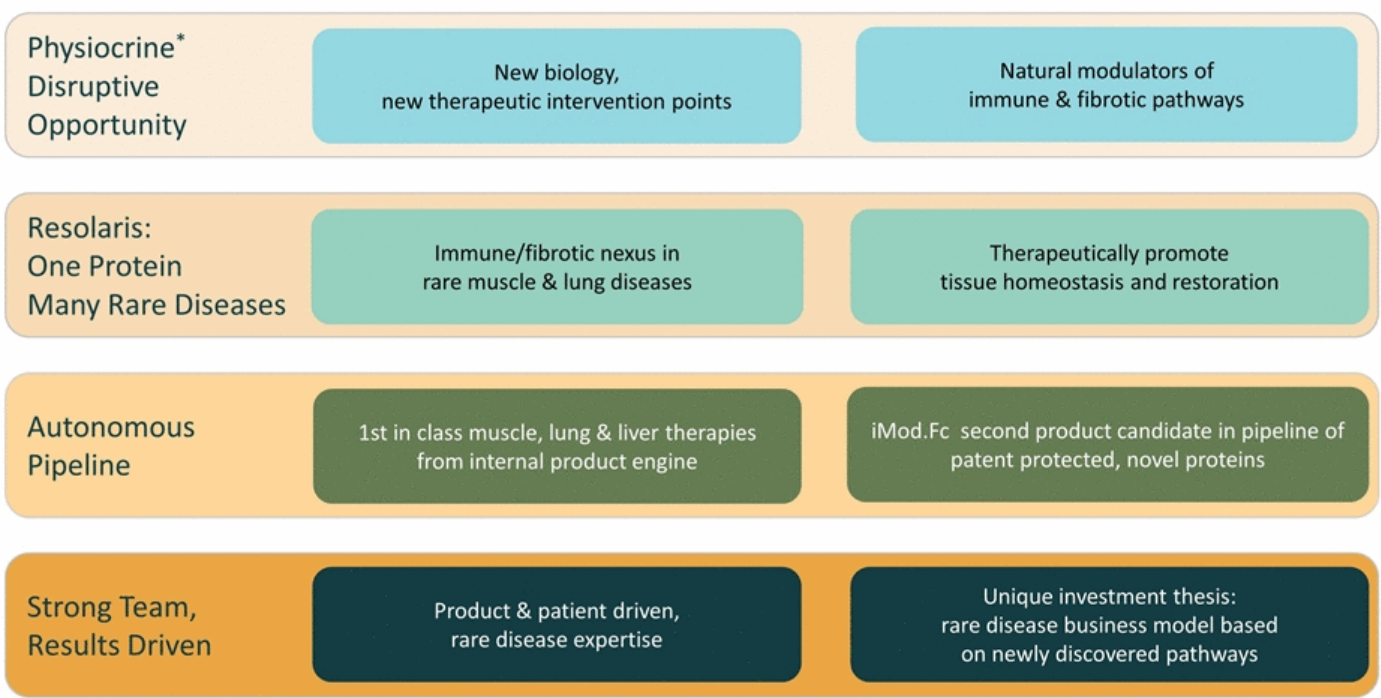
# Forward-Looking Statements

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The following slides and any accompanying oral presentation contain forward-looking statements within the meaning of the Private Securities Litigation Reform Act of 1995 and other federal securities laws. The use of words such as “may,” “might,” “will,” “should,” “expect,” “plan,” “anticipate,” “believe,” “estimate,” “project,” “intend,” “future,” “potential,” “opportunity,” or “continue,” and other similar expressions are intended to identify forward-looking statements. For example, all statements we make regarding the potential therapeutic benefits of Physiocrines and our product candidates, including Resolaris™ and iMod. Fc, the ability to successfully advance our pipeline or product candidates, the timing within which we expect to initiate, receive and report data from, and complete our planned clinical trials, and our ability to receive regulatory approvals for, and commercialize, our product candidates, our ability to identify and discover additional product candidates, and the ability of our intellectual property portfolio to provide protection are forward-looking statements. All forward-looking statements are based on estimates and assumptions by our management that, although we believe to be reasonable, are inherently uncertain. All forward-looking statements are subject to risks and uncertainties that may cause actual results to differ materially from those that we expected. These risks, uncertainties and other factors are more fully described in our filings with the U.S. Securities and Exchange Commission, including our most recent Quarterly Report on Form 10-Q. The forward-looking statements in this presentation speak only as of the date of this presentation and neither we nor any other person assume responsibility for the accuracy and completeness of any forward-looking statement. We undertake no obligation to publicly update or revise any forward-looking statement, whether as a result of new information, future events or otherwise, except as required by law.

We own various U.S. federal trademark applications and unregistered trademarks, including our company name and Resolaris™. All other trademarks or trade names referred to in this presentation are the property of their respective owners. Solely for convenience, the trademarks and trade names in this presentation are referred to without the symbols ® and ™, but such references should not be construed as any indicator that their respective owners will not assert, to the fullest extent under applicable law, their rights thereto.





\*Proteins for life (physio) specific activity (crine)

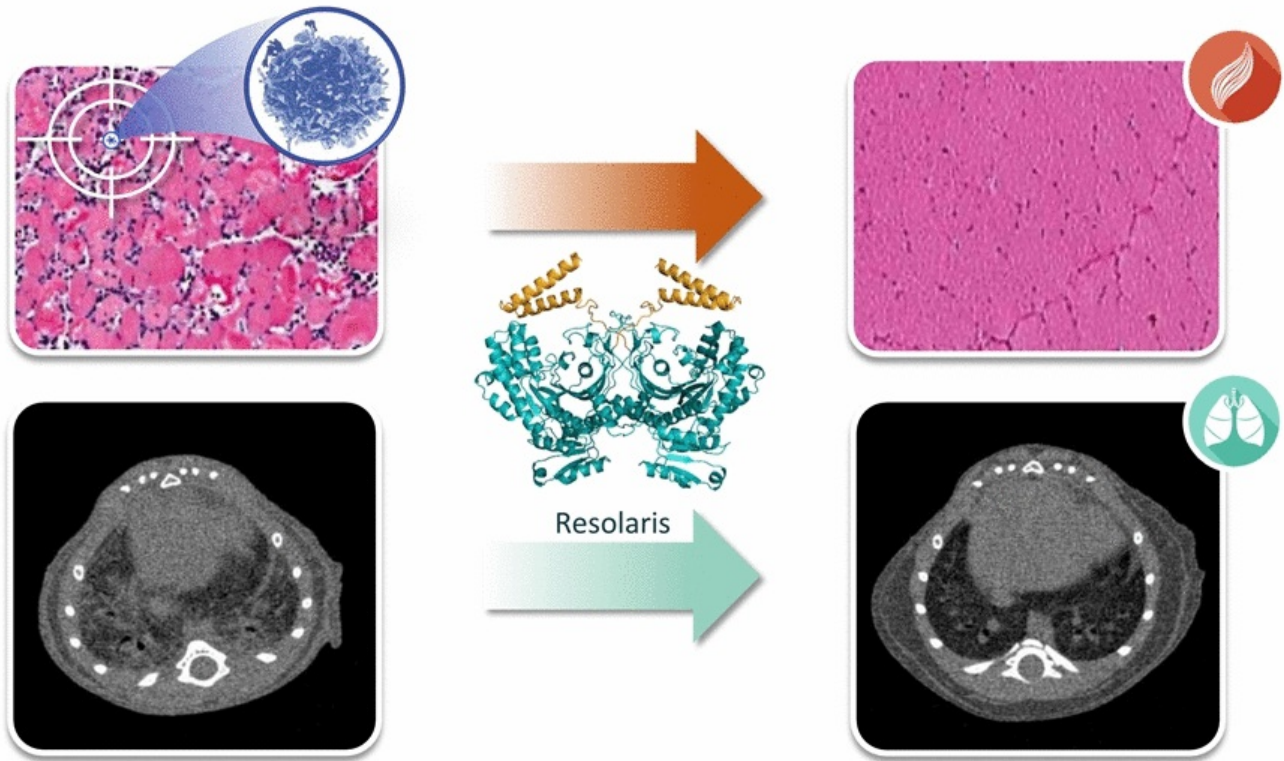




# Resolaris Improves Muscle & Lung Immune/Fibrotic Components

FIRST  
PROGRAM

*Two weeks of treatment in Statin myopathy & Bleomycin IPF\* models*



\*Idiopathic Pulmonary Fibrosis ("IPF"), a subset of Interstitial Lung Disease ("ILD")





# Immuno/Fibrosis Modulation Pipeline

PARADIGM SHIFT  
IN TREATMENTS

*1<sup>st</sup> in class candidates for rare diseases with an immune or fibrotic component*



**Patient Phenotype Focus**

- Severe impact from disease with the potential for large treatment effect
- Subject to poor standard of care

<sup>1</sup> Facioscapulohumeral Muscular Dystrophy  
<sup>2</sup> Limb-girdle Muscular Dystrophy  
<sup>3</sup> Interstitial Lung Disease

# Leadership Team

EXPERIENCED  
INDUSTRY VETERANS



**John Mendlein, Ph.D.**  
Chief Executive Officer



**Sanuj Ravindran, M.D.**  
Chief Business Officer



**Melissa Ashlock, M.D.**  
SVP, Translational Medicine  
& Therapeutics



**Kelly Blackburn**  
VP, Clinical Operations



**John McKew, Ph.D.**  
VP, Research



**Andrew Cubitt, Ph.D.**  
VP, Product Protection




**Ashraf Amanullah, Ph.D.**  
VP, Manufacturing



**John Blake, CPA**  
VP, Finance



A woman with dark hair and glasses, wearing a white hoodie and blue jeans, is seated in a wheelchair. She is looking towards the camera. The background is a digital-themed image with vertical columns of binary code (0s and 1s) in shades of blue and green. On the left side, there is a faint, glowing molecular or cellular structure. The overall color palette is cool, dominated by blues and greens.

***Discovery of Physiocrine Proteins:  
Alternative Splicing of Ancient Genes  
Leads to New Biology***





# Dual Roles of tRNA Synthetases: Protein Synthesis & Potential for Physiological Regulation

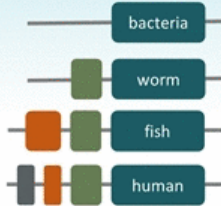
GENETIC BASIS OF  
PHYSIOCRINES

tRNA Synthetases:  
In all cellular life forms  
*4 billion years old*

Intracellular Function:  
23 genes in humans  
*Protein synthesis*

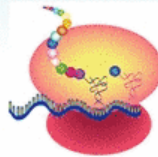
Extracellular Function:  
~300 Physiocrines  
*Splice variants offer new biology*

Physiocrine Regions of tRNA  
Synthetase displayed in a gene



Intracellular  
*(Original Primordial Function)*

Protein Synthesis (retained in all species)



Extracellular Physiological  
*(Systems Evolved with Species)*

Regenerative Systems (worm)

Vascular Systems (fish)

Immune Systems (human)

## Therapeutic Opportunities

Augment Homeostatic Pathways by Administering Physiocrine Proteins

Orchestrate multiple physiological processes

Focus on Immuno- & Fibro-modulation

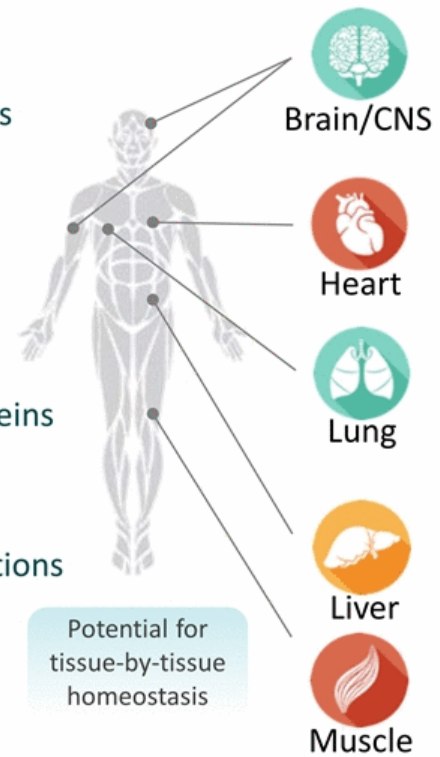


# Primer on Physiocrines

*Potential for novel physiological modulation in disease*

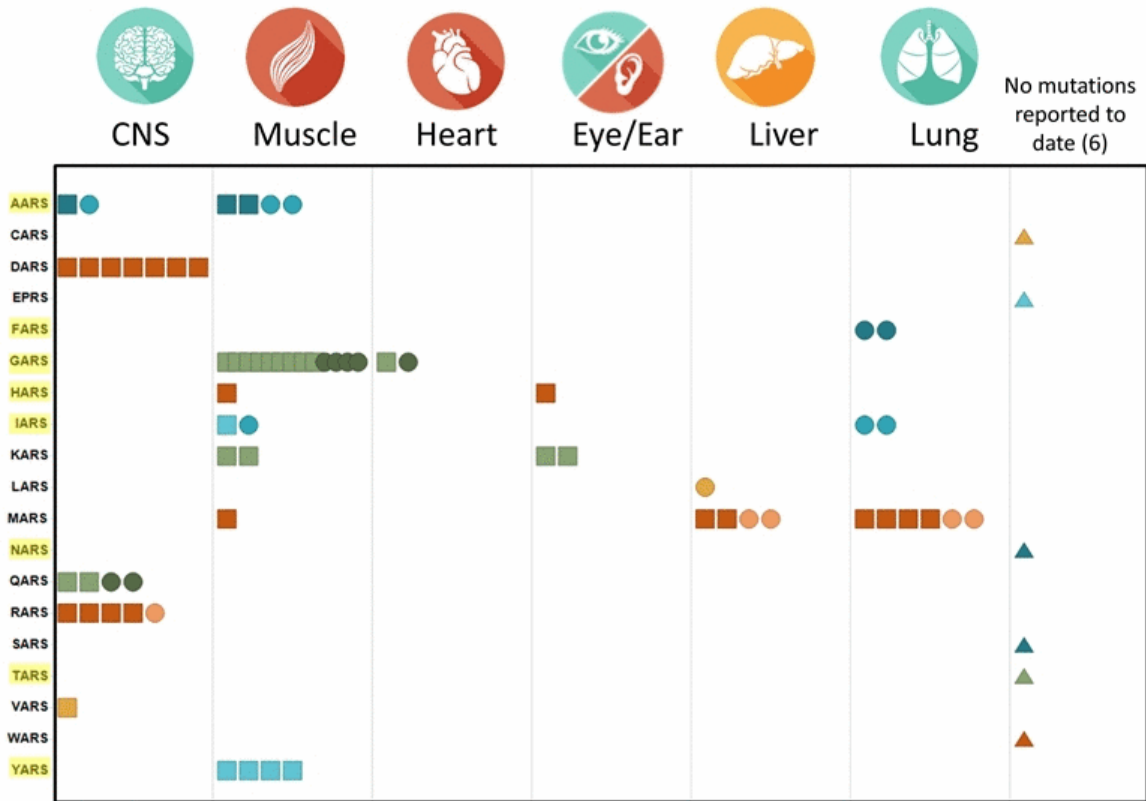
PHYSIOCRINE  
DISCOVERY

- Extracellular functions of intracellular tRNA synthetases
- New class of modulators of tissue homeostasis
- Conserved, genetic gain of function over time
- ~300 proteins varying in size (40-500AA)
- Work via GPCRs, TLRs, cytokine receptors & other proteins
- Not glycosylated & non-canonical leader sequences
- At least 14/23 members of family with disease connections



# Physiocrine Mutations in Disease Organ Phenotypes

GENETIC EVIDENCE OF THE IMPORTANCE OF PHYSIOCRINES



Domain location of mutation:

- Physiocrine
- tRNA synthetases targeted by autoantibodies
- Catalytic (may overlap with Physiocrine domain)



# Leading Journal Publications: Science & Nature



Published in Luo et al., Science, 2014; Weiwei, He, et al., Nature 2015. Graphics are visual representations for illustrative purposes.







***The First Physiocrine, the Resokine Pathway:  
Potential New Therapies  
in Rare Muscle & Lung Diseases***



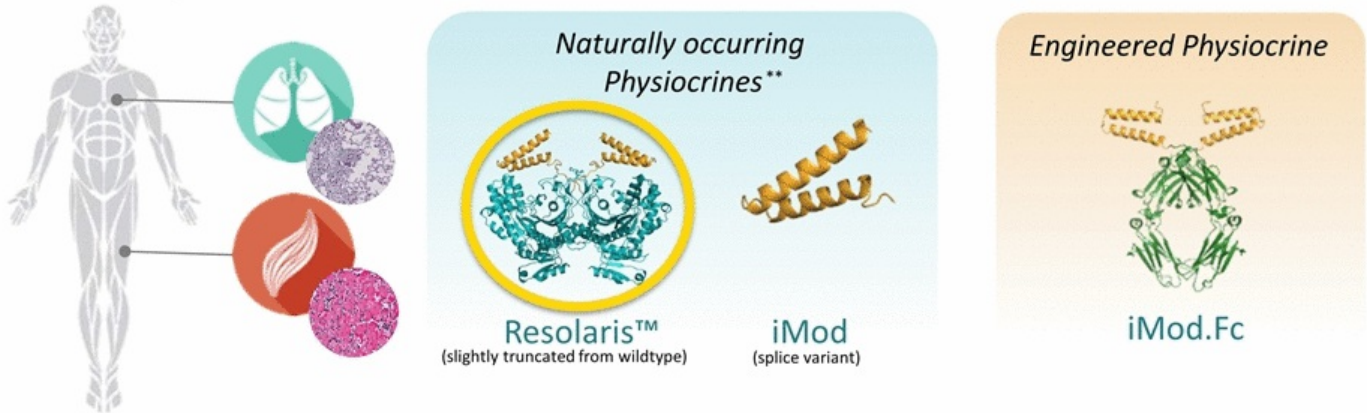
# Molecules of the Resokine Pathway

RESOKINE  
FRANCHISE



*New hope for rare muscle & lung disease patients*

## Resokine\* Pathway Discovery:

- Immuno-modulatory & fibro-modulatory activity via in vivo screening
- Clinical correlates with Jo-1 Ab phenotype: myopathy & ILD where we believe the pathway is disrupted



## Therapeutic applications with a Jo-1 Ab phenotype:

- Rare myopathies with an immune component (RMICs) 
- Rare pulmonopathies with an immune component (RPICs) 
  - Pursuing subsets of ILD

\*Resokine (name for extracellular HARS, a tRNA Synthetase)

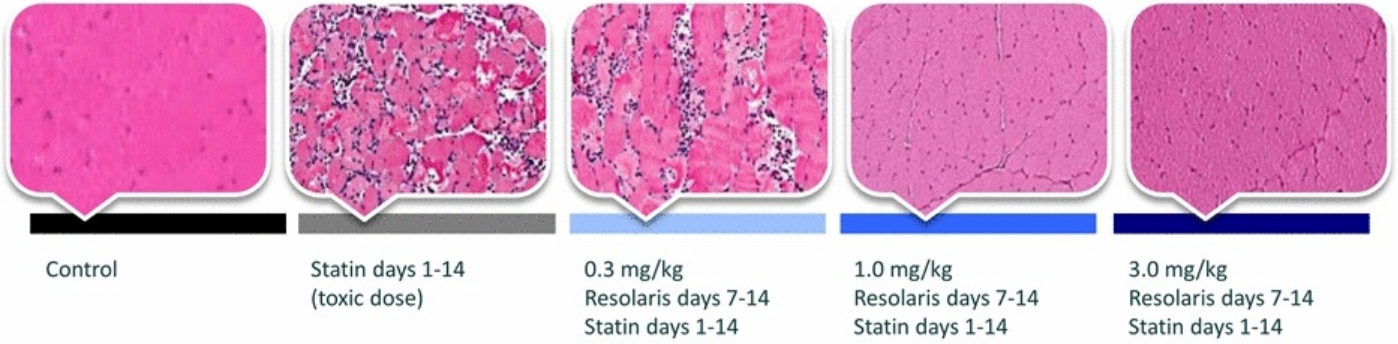
\*\*Zhou et al, JBC, 2014



# Ability to Treat Immune Cell Invasion in Muscle

Myopathy model: *in vivo* Resolaris effects

RESOLARIS  
PRECLINICAL



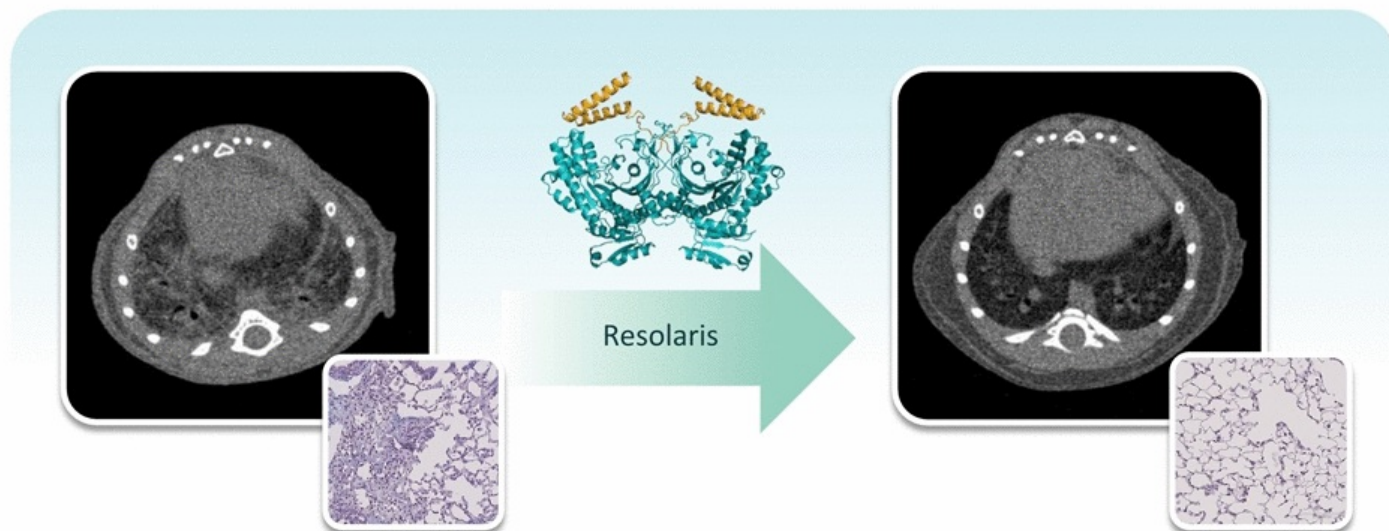
↓ Cytokines, ↓ T-cells and ↓ Monocytes with Resolaris administration




# Resolaris: Active In Lung Inflammation & Fibrosis Model

Three week rodent model, two weeks of therapeutic treatment

PROMISING  
THERAPEUTIC  
ACTIVITY



 Pulmonary Inflammation and Fibrosis Induced with Bleomycin  
Promising therapeutic activity\*  
Compared favorably to Pirfenidone

Experimental data provided by Stelic CRO  
CT scans taken at day 14, lung histology taken at day 21  
\* Activity of mouse Resolaris (3mg/kg) vs vehicle control

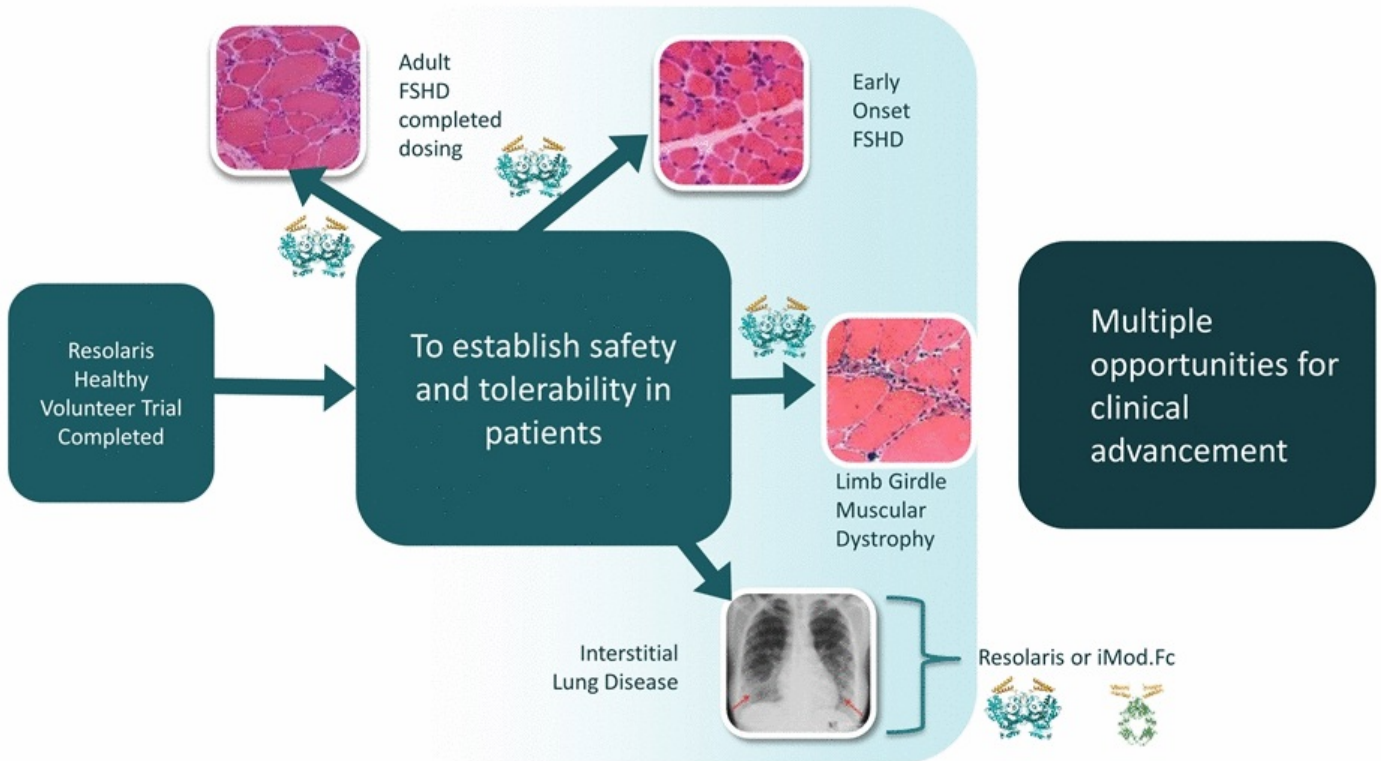


***Clinical Development of Resolaris:  
Harnessing a Natural Pathway  
to Treat Multiple Muscle & Lung Diseases***

# Clinical Path for Resolaris™ & iMod.Fc

Staging Rare Muscle & Lung Disease Indications

RESOLARIS  
CLINICAL





# Resolaris for Multiple Muscle Indications

Rare myopathies with an immune component (RPIC)

RESOLARIS  
CLINICAL

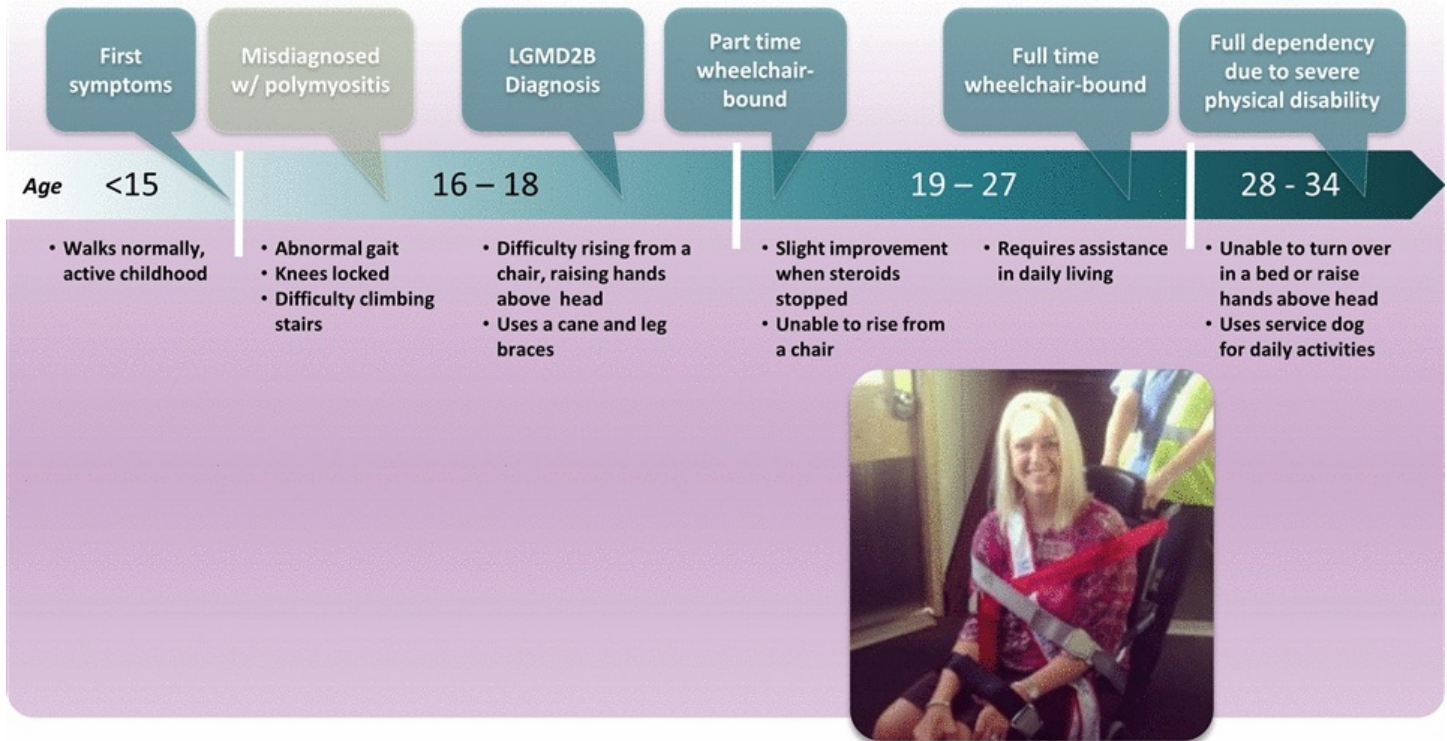
	Facioscapulohumeral Muscular Dystrophy (FSHD)	Limb-girdle Muscular Dystrophy 2B (LGMD2B)
Pathology	<ul style="list-style-type: none"><li>▪ Immune component (e.g. T cells)</li><li>▪ Toxic gain of function mutation (Dux4 protein)</li><li>▪ Muscle-by-muscle progression; MRI assessable</li></ul>	<ul style="list-style-type: none"><li>▪ Immune component (e.g. T-cells)</li><li>▪ Toxic loss of function mutation (dysferlin)</li><li>▪ Muscle group progression; MRI assessable</li></ul>
Clinical	<ul style="list-style-type: none"><li>▪ Debilitating muscle weakness</li><li>▪ Difficulty raising arms, foot drop when walking</li><li>▪ May have visual or auditory impairment</li></ul>	<ul style="list-style-type: none"><li>▪ Debilitating muscle weakness</li><li>▪ Challenges moving limbs</li><li>▪ May have respiratory insufficiency</li></ul>
Standard of care	<ul style="list-style-type: none"><li>▪ No therapeutic treatments</li><li>▪ Only supportive care provided</li></ul>	
Trials	<ul style="list-style-type: none"><li>▪ <b>Adult FSHD P1b/2 completed dosing in cohort 3</b></li><li>▪ <b>Early onset FSHD P1b/2 trial initiated 4Q15</b></li><li>▪ Explore safety, tolerability, and activity</li></ul>	<ul style="list-style-type: none"><li>▪ <b>P1b/2 trial initiated in 4Q15</b></li><li>▪ <b>First patients dosed</b></li><li>▪ Explore safety, tolerability, and activity</li></ul>

*Similar clinical phenotype to Jo-1 Ab disease as both are sometimes misdiagnosed as either polymyositis or Jo-1 Ab Phenotype*



# LGMD2B Disease Progression Case History

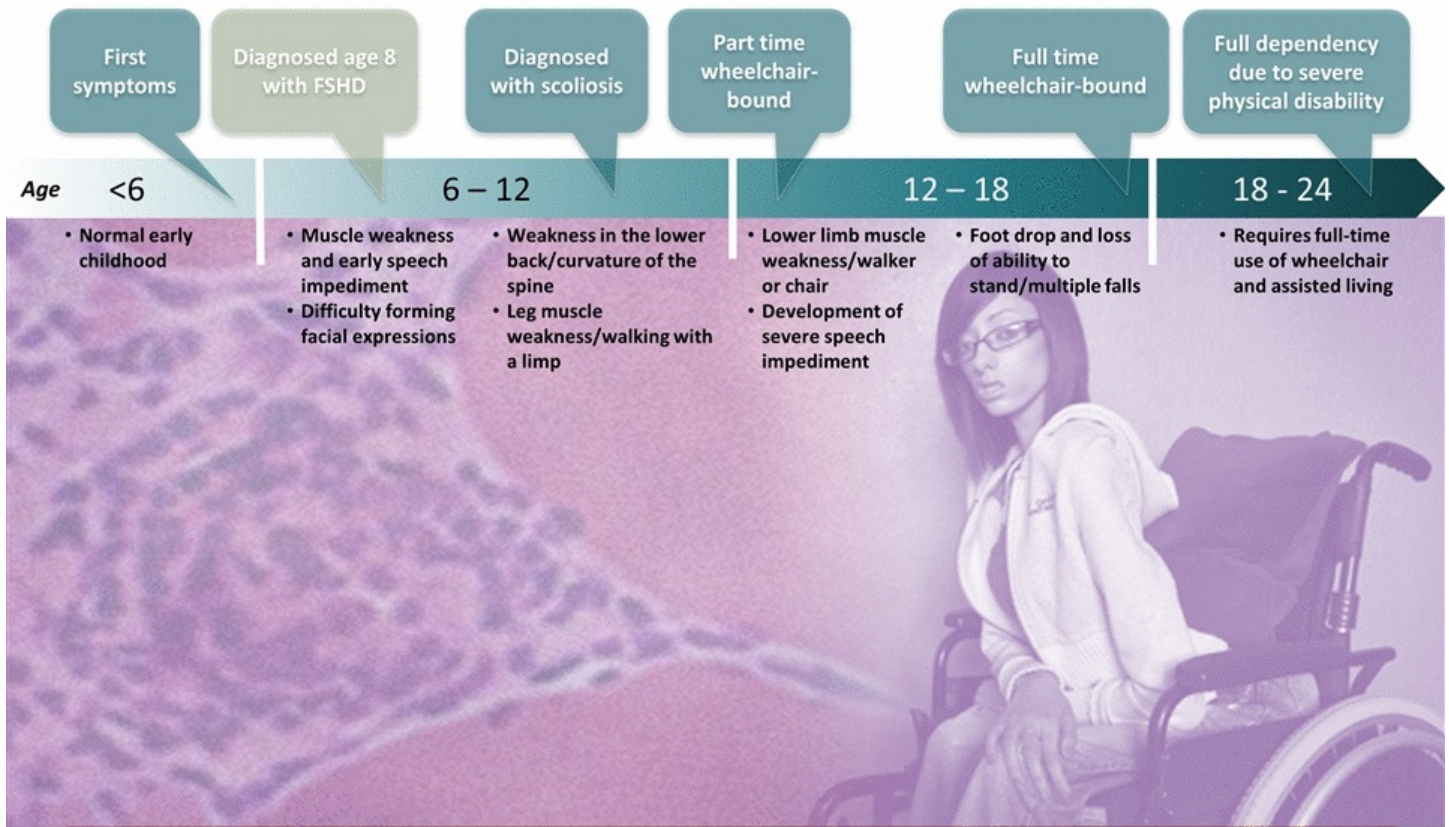
## Steroid-induced myopathy



<https://www.youtube.com/watch?v=JLqHis1yPUU>  
<http://mwtn2013blisswelch.blogspot.com/>



# Early Onset FSHD Case History

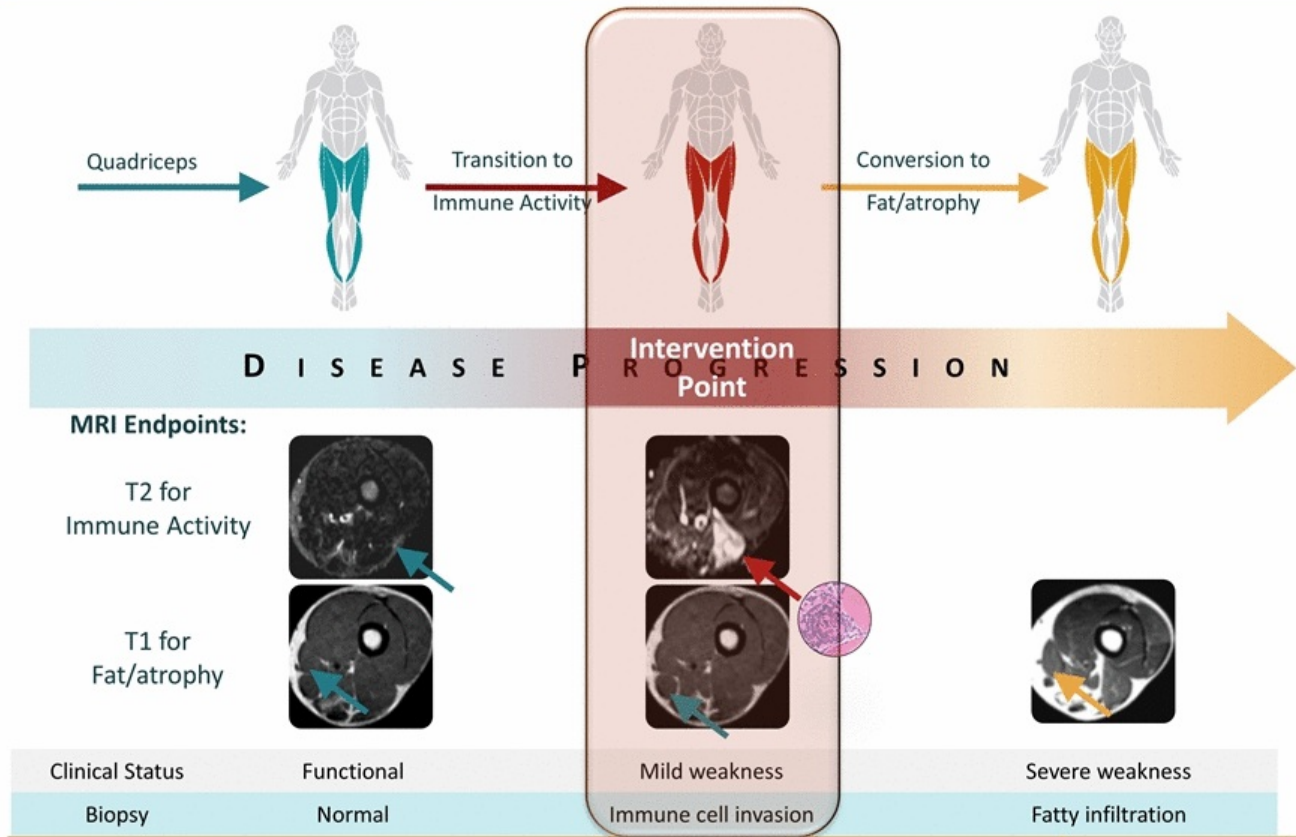


<http://www.theguardian.com/lifeandstyle/2009/may/28/muscular-dystrophy-disability-fshd>  
 Climbing Mountains; Sarabjit Parmar, 2014

# Monitoring Muscle-by-Muscle Disease Phenotype

Immunologically driven disease progression assessed by MRI

RESOLARIS  
CLINICAL



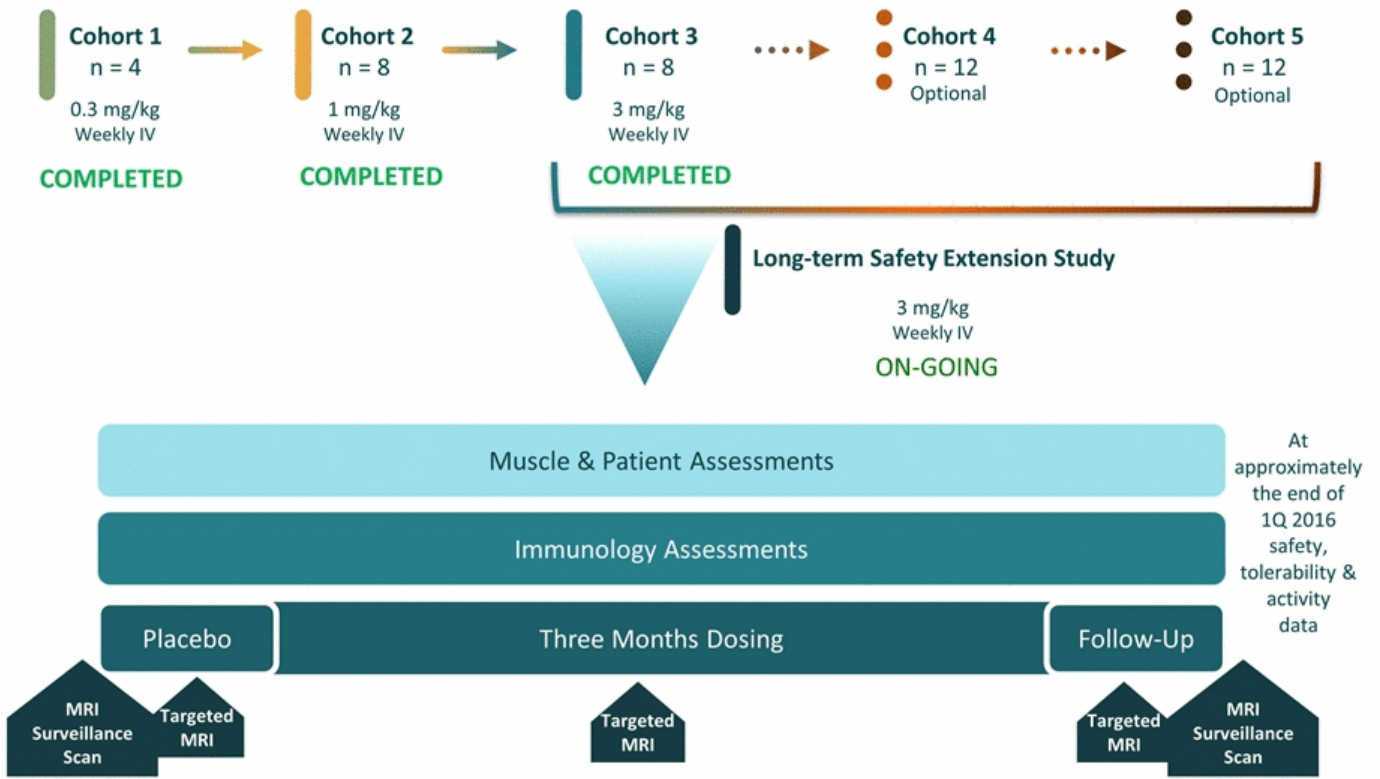
Frisullo et al., J Clin. Immunol., 2011; Also based on unpublished work and communications with investigators and KOLs



# Exploratory Study of Resolaris in Adult FSHD

Phase 1b/2 randomized, double-blind, placebo controlled, multiple ascending dose

RESOLARIS  
CLINICAL

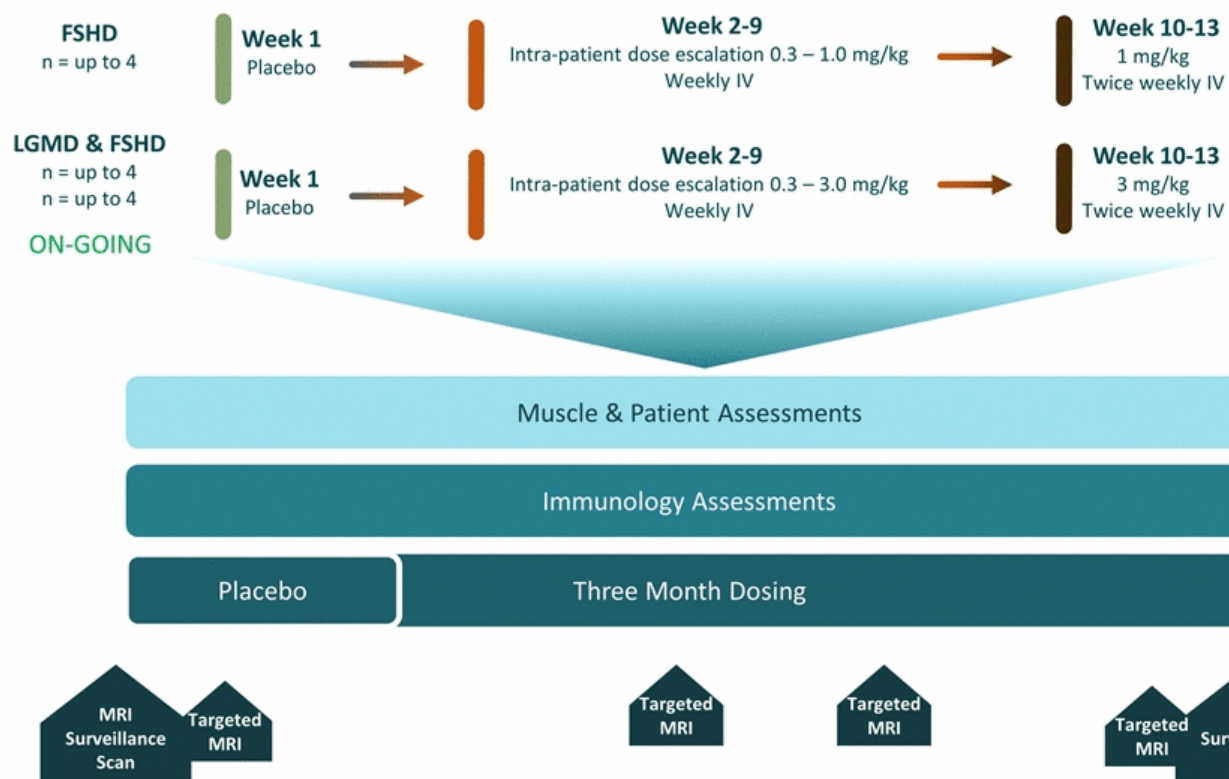




# LGMD2B/FSHD: Exploratory Study of Resolaris

Phase 1b/2 open-label, intra-patient dose escalation study

RESOLARIS  
CLINICAL



# Early Onset FSHD: Exploratory Study of Resolaris

RESOLARIS  
CLINICAL

Phase 1b/2 open-label, intra-patient dose escalation study

Early Onset FSHD ( $\leq 10$  as age of disease onset)

n = up to 16

**Stage 1**  
Ages 16-25  
n = up to 8



ON-GOING

Muscle & Patient Assessments

Immunology Assessments

Placebo | Three Month Dosing

MRI  
Surveillance  
Scan

MRI  
Surveillance  
Scan

**Stage 2**  
Ages 12-15  
n = up to 8



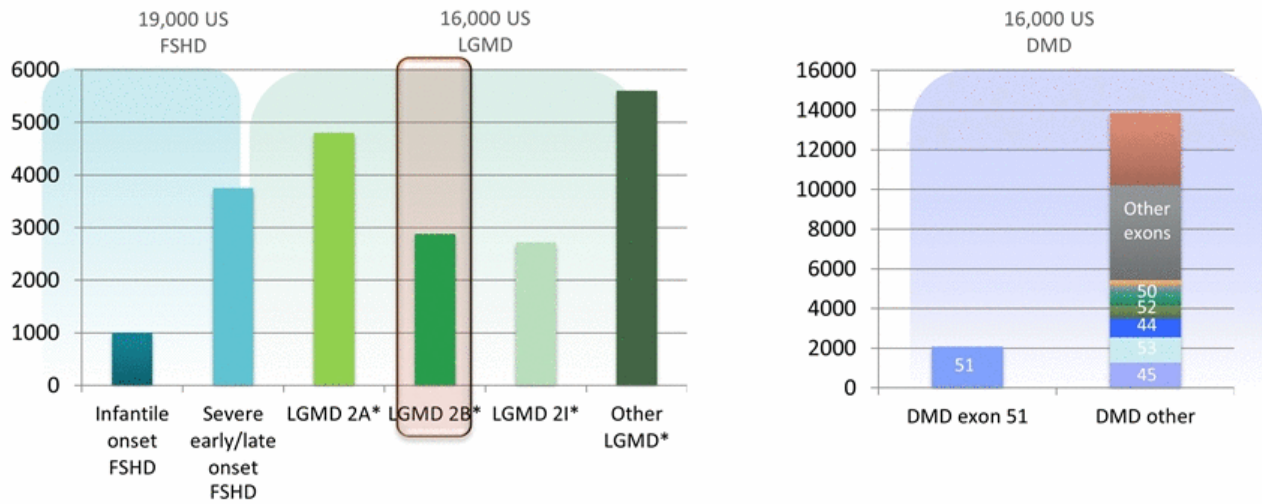
PLANNED



# Resolaris: One Product, Multiple Rare Diseases

Promise for grave, recalcitrant myopathy patients

COMMERCIAL



- Leverage registries, sites and advocacy groups to access FSHD/LGMD patients
- Leverage sites for indication expansion
- Common physician base for patient ID, prescribing and clinical development in RMICs

*FSHD: Average prevalence rates of FSHD are approximately 1/17,000. Applying this rate to the US population based on recent census data equals approximately 19,000.*  
*LGMD: 16,000 cases estimated in US population. 1/20,000 Wickland and Kissel, Neural. Clin. 2014. Relative Prevalence of Limb Girdle Muscular Dystrophies in the United States Population. Wicklund et al., Neurology 2013.*  
*DMD: Prevalence of approximately 5/100,000. Orphanet Report Series - Prevalence of rare diseases: Bibliographic data - May 2014 - Number 1*



## Interstitial Lung Disease > 80 RPIC Forms

### Pathogenesis

- Lung damage leading to alveolar inflammation or fibrosis
- Worst prognosis: lower DLCO and rapid decline of DLCO over three years

### Clinical manifestations

- Shortness of breath and cough
- Specific chest radiographic abnormalities
- Decreased lung volume noticed in pulmonary function tests

### Standard of care

- O<sub>2</sub>, pulmonary rehab
- Immunosuppressive (cyclophosphamide with low dose prednisone)
- Lung transplant

### Upcoming Trials

- **Expect to initiate exploratory P1b/2 trial with iMod.Fc (2017) and/or Resolaris (by the end of 2016)**
- Evaluating appropriate forms of ILD
- Goal is to explore safety, tolerability, biological and clinical activity



# An Engineered Physiocrine for Lung Disease: iMod.Fc

New TPP to open more lung indications

RESOKINE  
FRANCHISE

## Rationale for iMod.Fc\*:

- Resolaris TPP: Weekly dosing; limits lung applications
- Develop molecule for new TPP: potentially once-monthly dosing

## Product Concept:

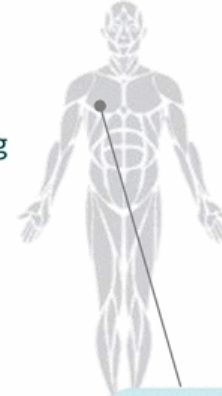
- Two iMod domains per Fc of an antibody
- Extend exposure to hit TPP

## Preclinical Path and Goals:

- Industry proven model of IPF (approved drugs: Pirfenidone & Nintedanib)
- *E. coli* produced for low COGs
- Show immuno- & fibro- modulatory activity
- Show safety at higher doses

## Potential Therapeutic applications:

- Rare pulmonopathies with an immune component (RPICs)
- Broader reach into RPICs and interstitial lung disease (ILD) indications



Rare Pulmonopathies with an Immune Component ("RPICs")



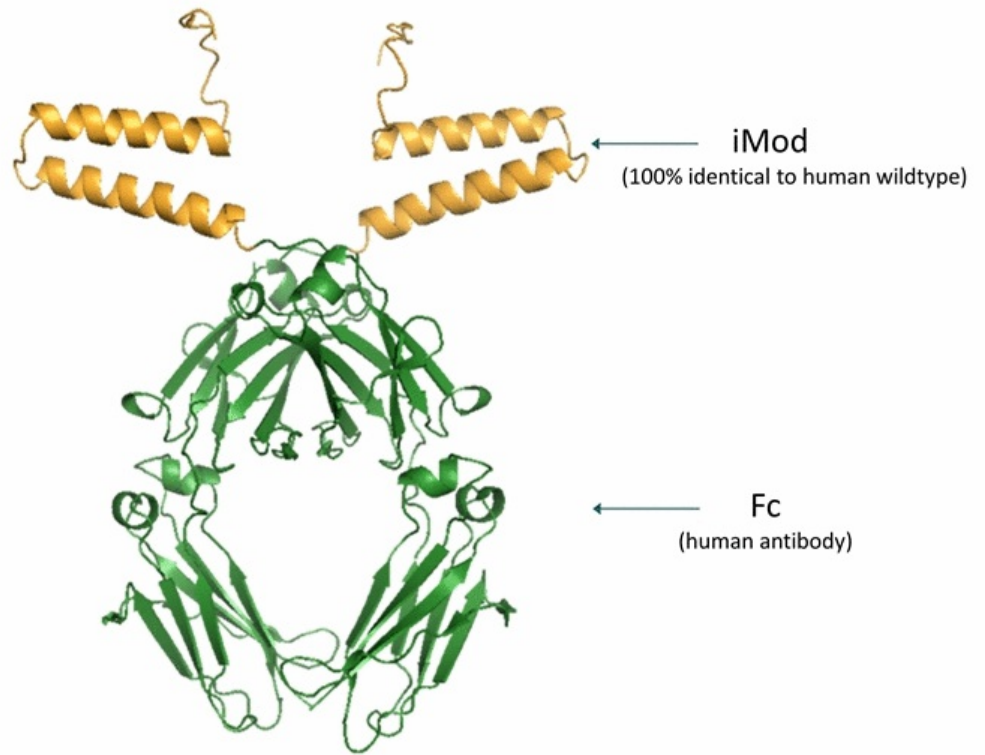
Lung

\* iMod.Fc refers to immunomodulatory domain of HARS fused to an Fc region of an antibody  
TPP = Target Product Profile



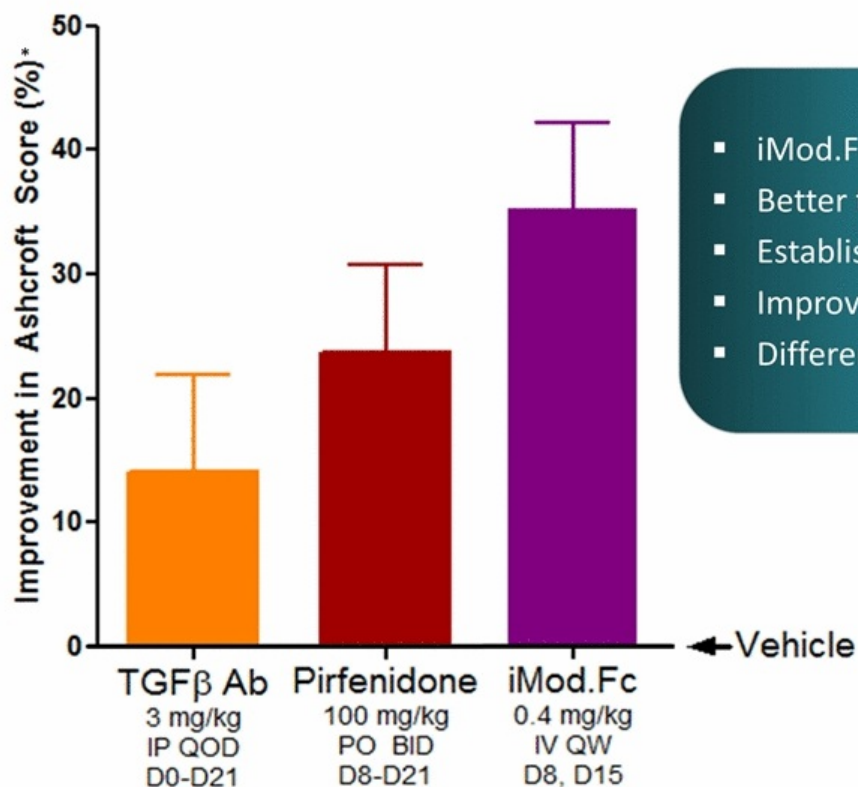
# iMod.Fc

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# Two iMod.Fc Doses Outperform 28 Pirfenidone Doses

SECOND  
IND CANDIDATE

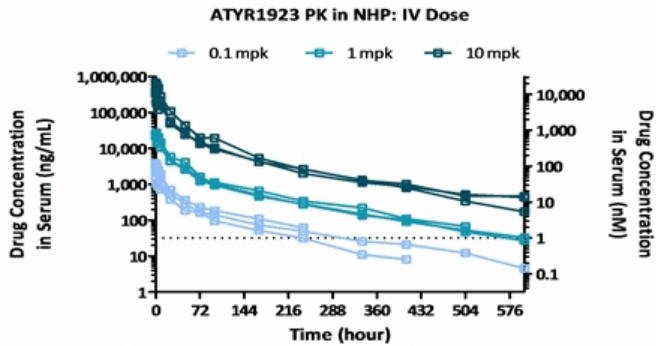


- iMod.Fc 1/250<sup>th</sup> of Pirfenidone dose
- Better than 10 TGFβ Ab doses
- Established IPF rodent model
- Improves inflammation & fibrosis
- Differentiated mechanism

\*The Ashcroft scale for the evaluation of bleomycin-induced lung fibrosis is the analysis of stained histological samples by visual assessment

# iMod.Fc: Non-Human Primate and Rodent PK and Safety

## Non-Human Primates



### Non-GLP double-dose study

- No clinical observations
- No pro-inflammatory cytokine signal

### Attractive PK

## Rodents

### Non-GLP toxicology

- 1-month study at dose level 25x efficacious dose
- No pro-inflammatory cytokine signal
- No clinical observations
- No changes in body or tissue weights
- 14 tissues examined microscopically; all normal

### Attractive PK

Supports potential for monthly dosing in patients







***Building a New Class of Therapeutics:  
Foundation for the Future***



## Strategic Assets for Programs and Class

- ~300 Physiocrines discovered
- Over 200 Physiocrines interrogated via protein expression and functional assays
- >200 pending applications worldwide
- 70 issued or allowed patents

## Global Patent Estate

- Claims to therapeutic compositions to block competitors
- Therapeutic compositions and methods of use including full length proteins
- Antibodies with selective binding specificity to a Physiocrine

## Resolaris Program

- US Patent protection to 2033, without patent extensions
- Composition of matter patents issued or allowed in the US and Europe
- Comprised of a number of patent families including U.S. Patent No. 8,835,387 covering Resolaris

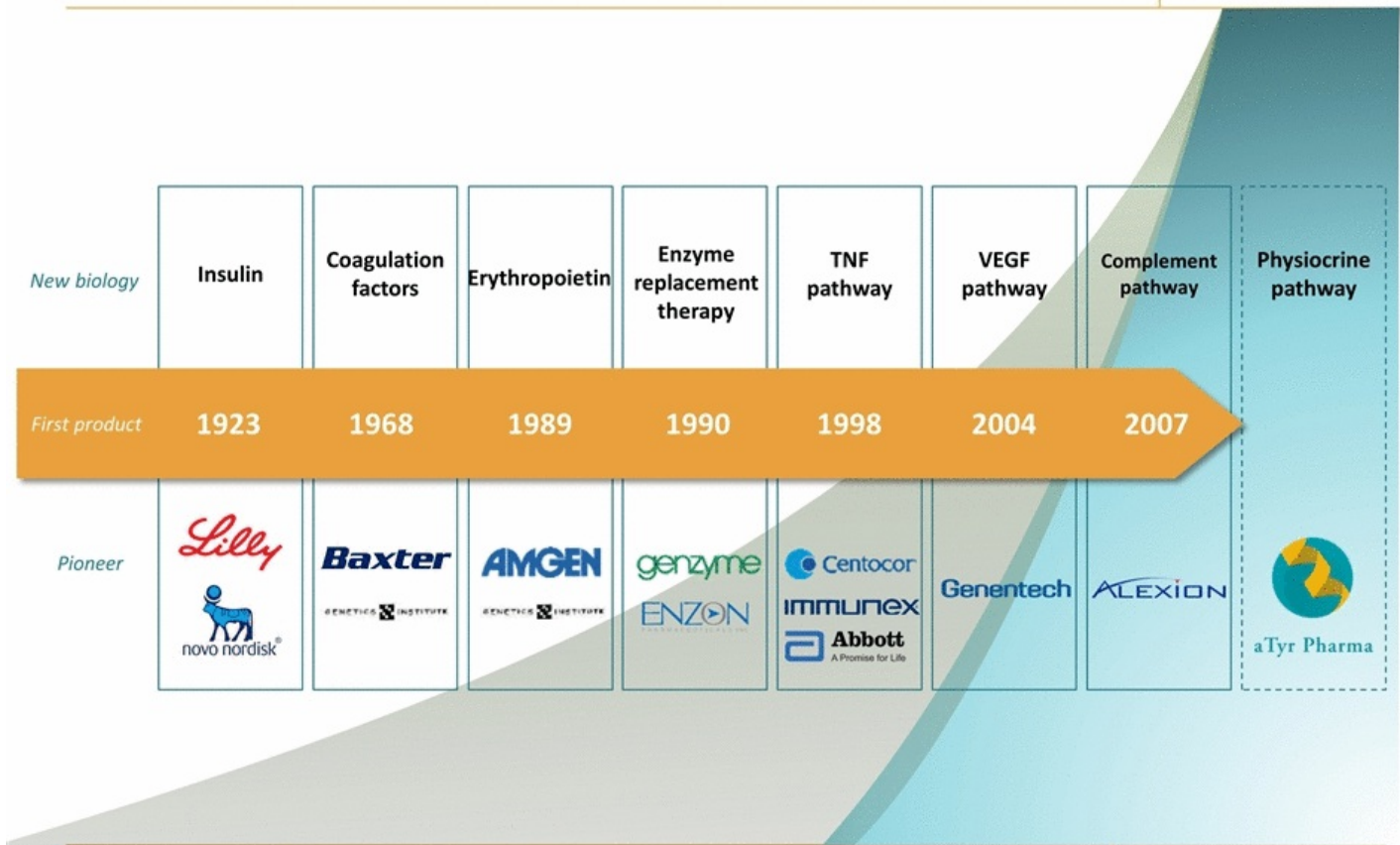




# Revolutionary Drugs Leveraging New Biology

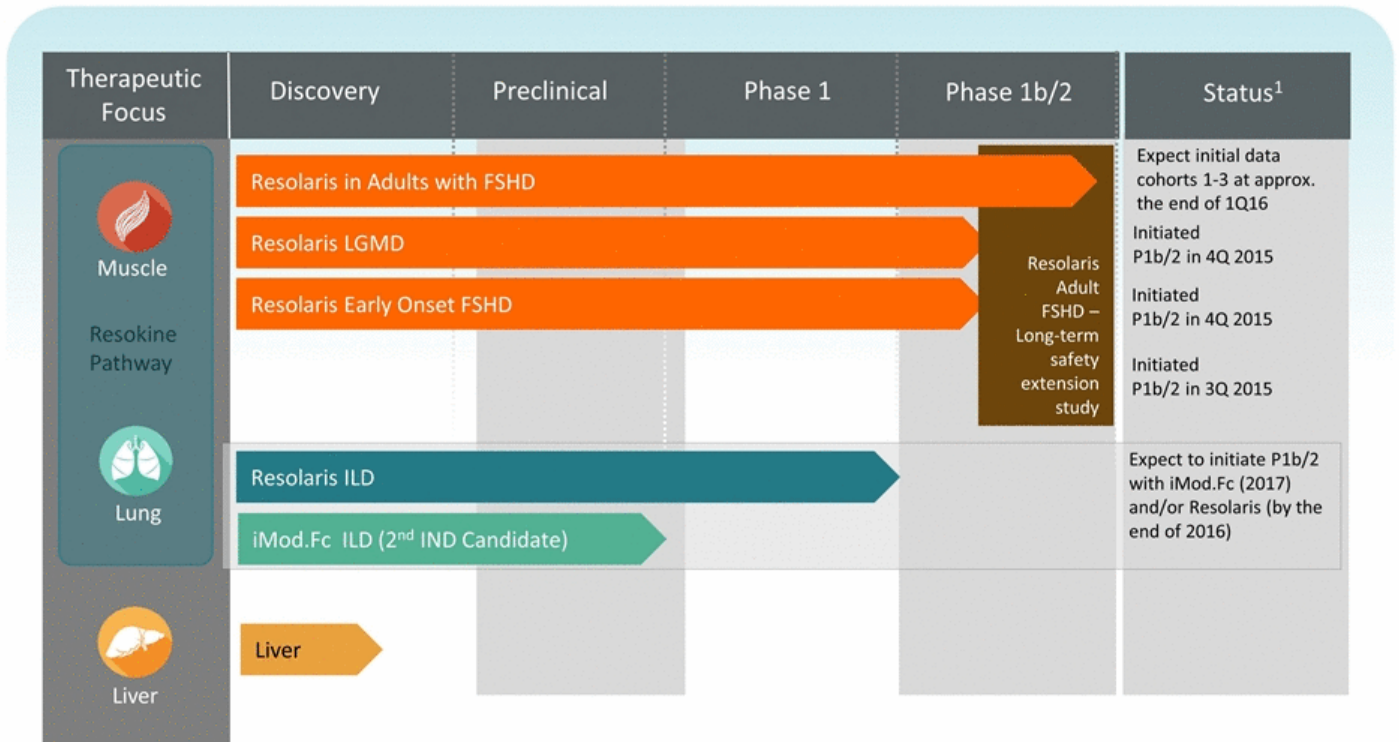
Opportunity to own a new class of meaningful medicines

MEANINGFUL  
MEDICINES



# Immuno/Fibrosis Modulation Pipeline

POTENTIAL  
FIRST-IN-CLASS  
MEDICINES



<sup>1</sup> The expected timing of the anticipated next milestones for our clinical programs for Resolaris in FSHD, LGMD and RPIC is based on our current estimates and is subject to change based upon a variety of risk factors.





**KEEP  
CALM  
AND  
MAINTAIN  
HOMEOSTASIS**

*Thank You*



