



**4th Quarter and Full Year
2021 Financial Results and
Update on Clinical
Development Programs**

February 28, 2022

Forward-Looking Statements

This presentation contains certain “forward-looking” statements that are made pursuant to the safe harbor provisions of the Private Securities Litigation Reform Act of 1995. All statements, other than statements of historical or present facts, are forward-looking statements, including statements regarding our future financial condition, future revenues, projected costs, prospects, business strategy, and plans and objectives of management for future operations, including our plans to submit for regulatory filings. In some cases, you can identify forward-looking statements by terminology such as “believe,” “will,” “may,” “might,” “estimate,” “continue,” “anticipate,” “intend,” “target,” “project,” “model,” “should,” “would,” “plan,” “expect,” “predict,” “could,” “seek,” “goal,” “potential,” or the negative of these terms or other similar terms or expressions that concern our expectations, strategy, plans, or intentions. These statements are based on our intentions, beliefs, projections, outlook, analyses, or current expectations using currently available information, and are not guarantees of future performance, and involve certain risks and uncertainties. Although we believe that the expectations reflected in these forward-looking statements are reasonable, we cannot assure you that our expectations will prove to be correct. Therefore, actual outcomes and results could materially differ from what is expressed, implied, or forecasted in these statements. Any differences could be caused by a number of factors including but not limited to: our expectations regarding the timing, costs, conduct, and outcome of our clinical trials, including statements regarding the timing of the initiation and availability of data from such trials; the timing and likelihood of regulatory filings and approvals for our product candidates; whether regulatory authorities determine that additional trials or data are necessary in order to obtain approval; our ability to obtain funding for our operations, including funding necessary to complete further development and commercialization of our product candidates; our plans to research, develop, and commercialize our product candidates; the commercialization of our product candidates, if approved; the rate and degree of market acceptance of our product candidates; our expectations regarding the potential market size and the size of the patient populations for our product candidates, if approved for commercial use, and the potential market opportunities for commercializing our product candidates; the success of competing therapies that are or may become available; our expectations regarding our ability to obtain and maintain intellectual property protection for our product candidates; the ability to license additional intellectual property relating to our product candidates and to comply with our existing license agreements; our ability to maintain and establish relationships with third parties, such as contract research organizations, contract manufacturing organizations, suppliers, and distributors; our ability to maintain and establish collaborators with development, regulatory, and commercialization expertise; our ability to attract and retain key scientific or management personnel; our ability to grow our organization and increase the size of our facilities to meet our anticipated growth; the accuracy of our estimates regarding expenses, future revenue, capital requirements, and needs for additional financing; our expectations related to the use of our available cash; our ability to develop, acquire, and advance product candidates into, and successfully complete, clinical trials; the initiation, timing, progress, and results of future preclinical studies and developments and projections relating to our competitors and our industry; the impact of governmental laws and regulations and regulatory development in the United States and foreign countries; the impact of the coronavirus disease (COVID-19) on our clinical trials, our supply chain, and our operations; and other risks and uncertainties, including those described under the heading “Risk Factors” included in our most recent Annual Report on Form 10-k for the year ended December 31, 2021, filed with the U.S. Securities and Exchange Commission (SEC) on February 28, 2022.

Additional factors that could cause actual results to differ materially from our expectations can be found in our Securities and Exchange Commission filings. Moreover, we operate in a very competitive and rapidly changing environment. New risk factors emerge from time to time, and it is not possible for our management to predict all risk factors, nor can we assess the effects of all factors on our business or the extent to which any factor, or combination of factors, may cause actual results to differ materially from those contained in, or implied by, any forward-looking statements. All forward-looking statements included in this presentation are expressly qualified in their entirety by these cautionary statements. The forward-looking statements speak only as of the date made and, other than as required by law, we undertake no obligation to publicly update or revise any forward-looking statements, whether as a result of new information, future events, or otherwise.

Omaveloxolone, bardoxolone methyl, and RTA 901 are investigational drugs, and their safety and efficacy have not been established by any agency.

Today's Agenda

Opening Remarks | Warren Huff, CEO

Development Update | Colin Meyer, MD, Chief Innovation Officer and
Seemi Khan, MD, Chief Medical Officer

Financial & Operational Update | Manmeet Soni, President

Concluding Remarks | Warren Huff, CEO

Regulatory Update on Omaveloxolone for Patients with Friedreich's Ataxia



Obtained **Fast Track Designation**



Initiated **rolling submission of NDA**



Plan to complete NDA submission by **end of 1Q 2022**



Update on Bardoxolone Program



Received CRL from FDA for **bardoxolone in Alport syndrome**



We will continue to work with the FDA to **confirm our next steps on our Alport syndrome program**



FALCON protocol amendment submitted to FDA
Type A meeting requested



Omaveloxolone in
Friedreich's
Ataxia

 MOXIe
a study in Friedreich's ataxia

 REATA.
PHARMACEUTICALS

Friedreich's Ataxia: No Approved Therapy



Patients progressively lose motor function

Typically diagnosed in teens,¹ wheelchair-bound in twenties²



Mean survival in mid-thirties³



Caused by epigenetic silencing of frataxin

Impaired mitochondrial function, suppressed Nrf2 expression, which impairs energy production



Most common recessive form of ataxia

In U.S., an estimated 4,000 patients are diagnosed with Friedreich's ataxia⁴ out of 5,000 total⁵

¹Rummev C, et.al. *Neurol Genet.* 2019; ²Rummev C, et.al. *E Clinical Medicine.* 2020; ³Tsou AY, et. al. *J Neurol Sci.* 2011; ⁴2020 IQVIA claims data and projected diagnosed; ⁵Estimated by Reata and the Friedreich's Ataxia Research Alliance (FARA) based on publicly available data and from Vankan P, *J Neurochem* 2013

MOXIe Part 2: Summary of Efficacy

One of the largest completed global interventional studies in patients with FA

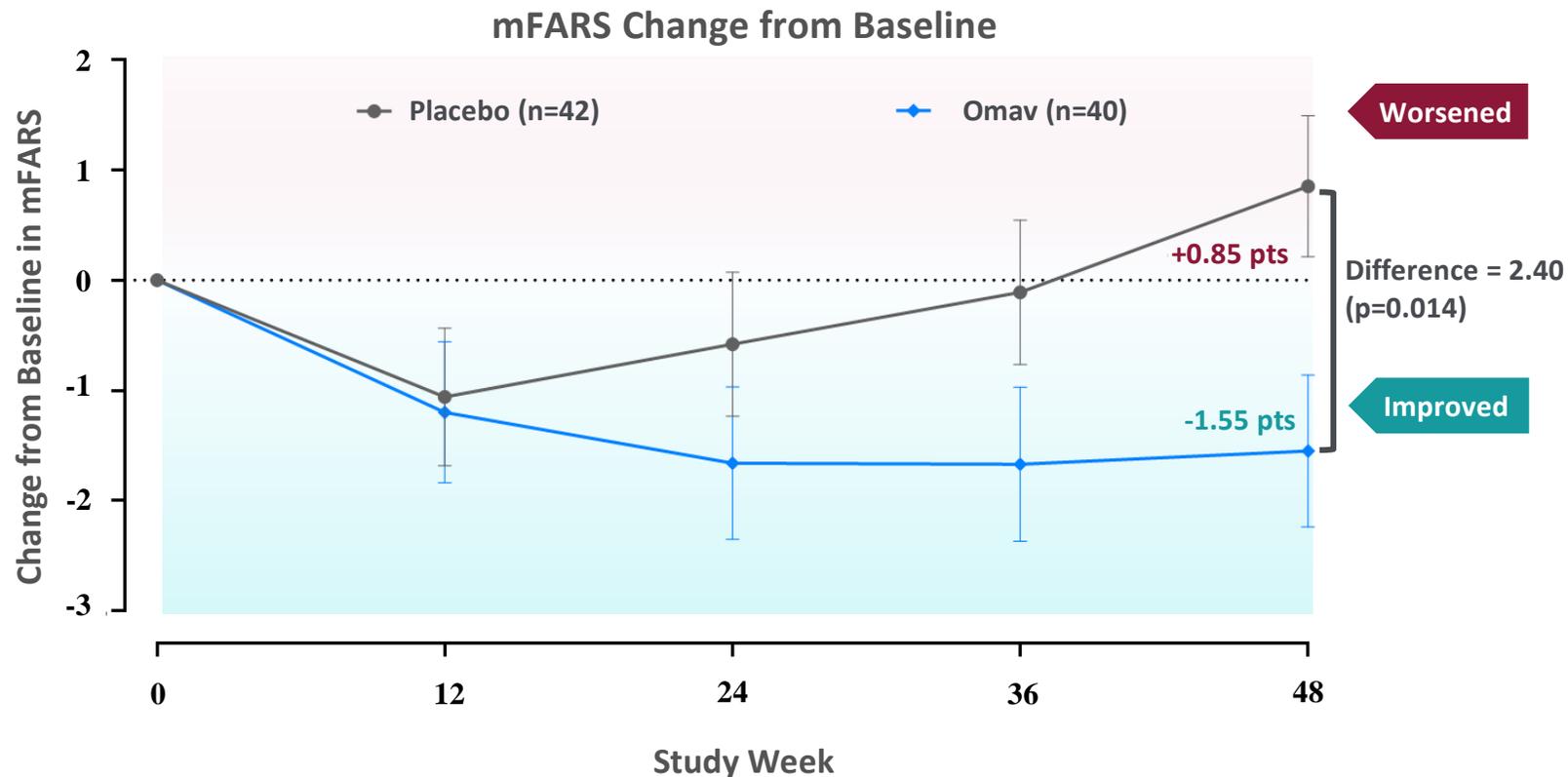
Enrolled a wide and representative range of patients with FA

- Baseline mFARS: 20 to 80
- Age: 16 to 40 years old

MOXIe Part 2 met its mFARS primary endpoint

- Improvement observed in all subsections relative to placebo
- Major subgroups and all analysis populations favored omaveloxolone

Improvement in several other efficacy measures assessed as secondary endpoints



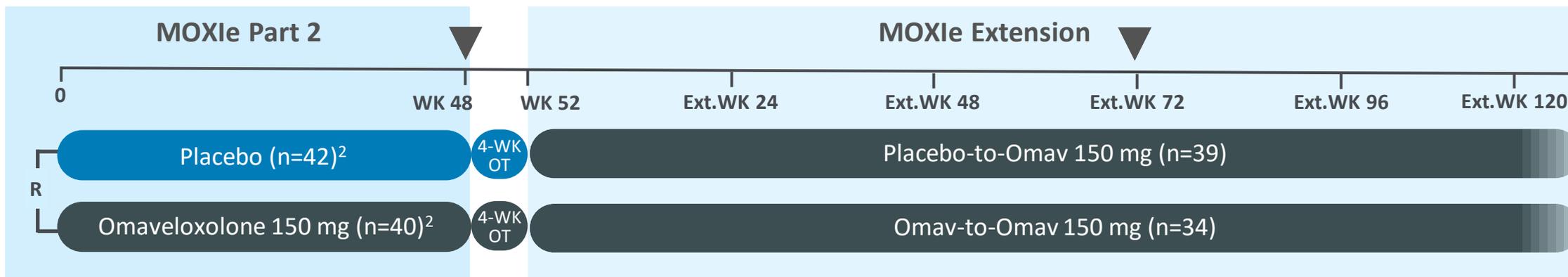
MOXle Extension: Delayed-Start Analysis Study Design

FDA suggested an exploratory, post-hoc analysis, the Delayed Start Analysis

- Established analysis methodology used in other neurological indications¹
- MOXle Extension is ongoing to continue evaluating efficacy and safety of omaveloxolone in patients with FA

Comparison of change from baseline in mFARS for patients without pes cavus randomized to placebo during MOXle Part 2 (Placebo-to-Omav) to patients randomized to omaveloxolone during MOXle Part 2 (Omav-to-Omav)

If the treatment effect at MOXle Extension Week 72 is maintained or non-inferior to the treatment effect at MOXle Part 2 Week 48, it demonstrates evidence of a persistent effect on disease course



¹Liu-Seifert, 2015a and 2015b; ²Patients without pes cavus and with a post-baseline mFARS assessment in MOXle Part 2; R: Randomization; OT: Off-treatment; WK: Week; Ext. WK: Extension week

Delayed-Start Analysis: Results Consistent with Persistent Effect on Disease Course

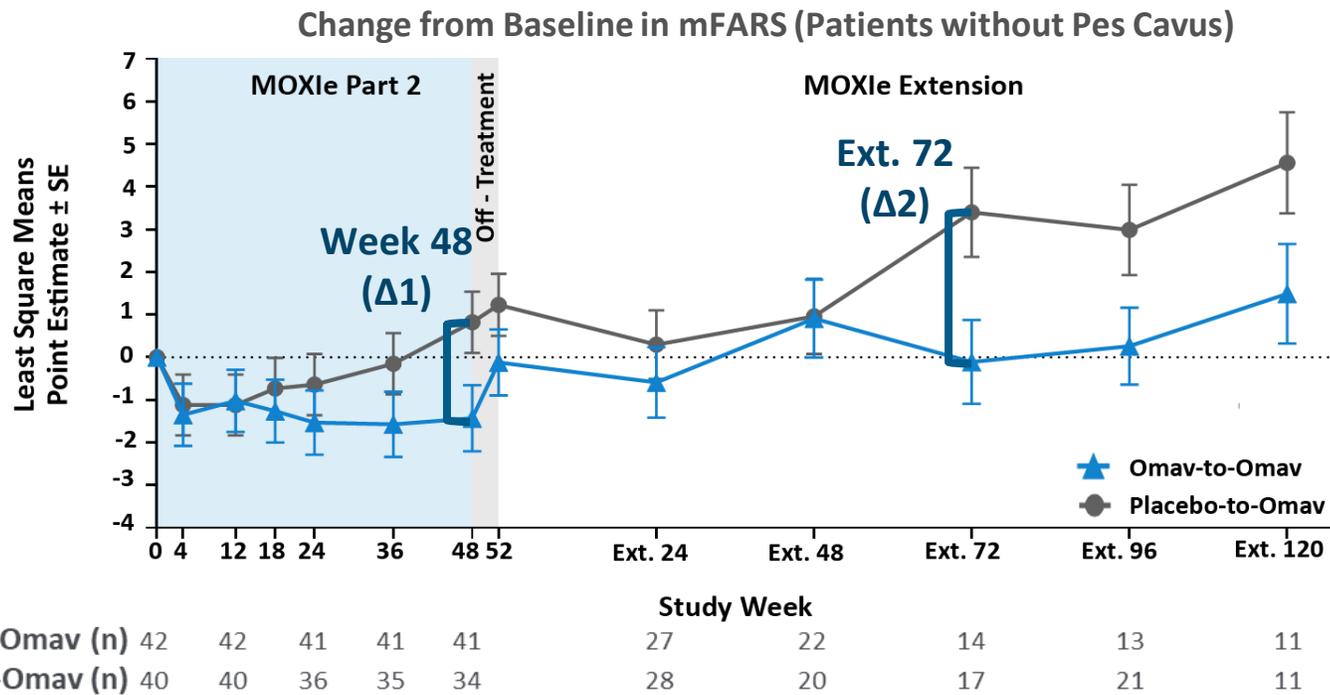
Non-inferiority test criteria met

- More than 50% of the difference between groups in mFARS observed at MOXle Part 2 Week 48 (-2.25, p=0.037) was preserved at MOXle Extension Week 72 (-3.51, p=0.016)
- Point estimate (-2.39) and upper limit of the 90% confidence interval (-0.615) less than zero

Delayed-Start Analysis (Non-Inferiority Test)¹

	MOXle Part 2 Week 48 ($\Delta 1$)	MOXle Extension Week Ext. 72 ($\Delta 2$)
Difference (LS Mean \pm SE)	-2.25 \pm 1.07 p=0.037	-3.51 \pm 1.45 p=0.016
Estimate = $\Delta 2 - 0.5 \times \Delta 1$	-2.39 \pm 1.38	
Upper Limit of 1-sided 90% CI for Estimate	-0.615	

¹Non-inferiority test performed using a mixed-model repeated measures analysis with a Toeplitz covariance structure



Delayed-Start Analysis: Parallel Annualized Slopes Consistent with Persistent Effect on Disease Course

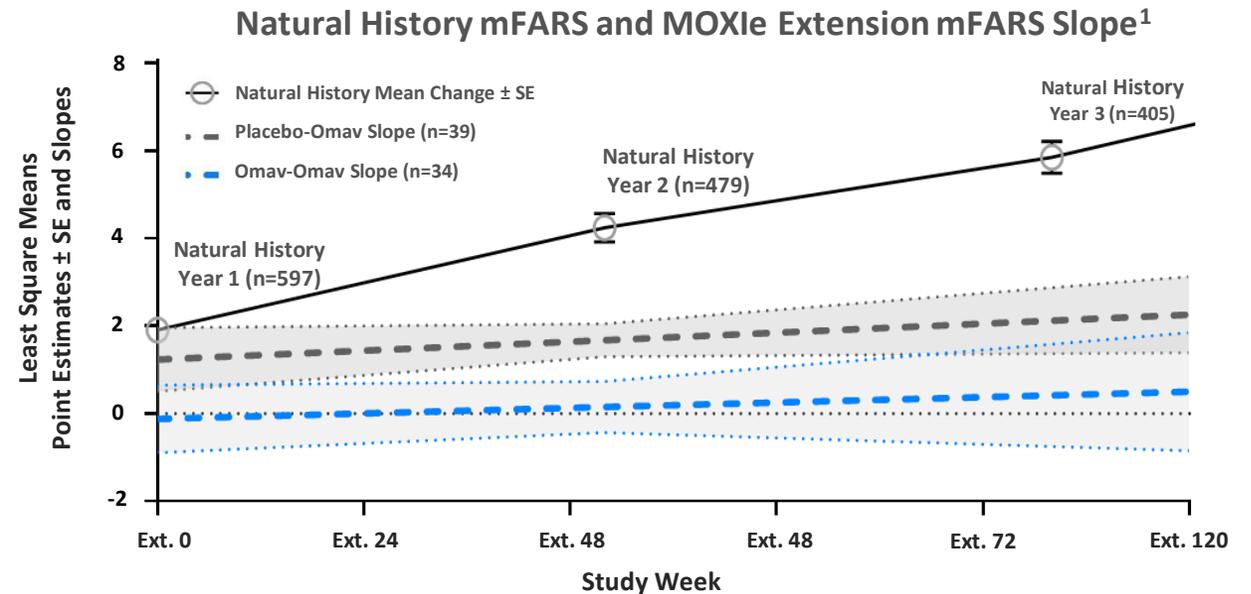
Longitudinal analysis was used to calculate annualized slopes in MOXle Extension

- Incorporates all available data from the MOXle Extension as of August 2021 data cut-off
- Placebo-to-Omav (0.45 ± 0.38) and Omav-to-Omav (0.27 ± 0.59)
- No significant difference between slopes (-0.18 ; $p=0.79$)

In MOXle Extension, omeveloxolone-treated patients have been progressing at a rate that is >75% less than the approximately two points per year that patients progressed in a recent large natural history study¹

Overall, the Delayed-Start Analysis results indicate a persistent omeveloxolone treatment effect on the course of FA

MOXle Extension Annualized mFARS Slope (\pm SE)		
Omav-to-Omav (n=34)	Placebo-to-Omav (n=39)	Difference
0.27 ± 0.59	0.45 ± 0.38	-0.18 ± 0.67 $p=0.79$



¹A mixed model repeated measures was used to fit change from baseline mFARS using all available data from MOXle Extension through Extension Week 120 to estimate annualized slopes based on MOXle Part 2 randomized treatment group. The slopes were calculated using a linear model with time, treatment, and the interaction of treatment and time as fixed factors. Natural History mean change \pm SE per year is based on Patel et. al. Ann. Clin. Transl. Neurol. 2016; Results based on August 2021 data cut-off.

MOXIe Part 2: Summary of Adverse Events

AEs generally mild to moderate in severity

- ALT and AST increases may be related to the pharmacological activity of omaveloxolone
- Not associated with other indicators of liver injury

SAEs

- SAEs reported in five omaveloxolone-treated patients (10%) and three placebo-treated patients (6%)

MOXIe Part 2 Summary of Adverse Events^{1,2}

Adverse Event	Placebo (n=52)	Omaveloxolone (n=51)
Headache	13 (25%)	19 (37%)
ALT increased	1 (2%)	19 (37%)
Nausea	7 (14%)	17 (33%)
Fatigue	7 (14%)	11 (22%)
Abdominal pain	3 (6%)	11 (22%)
AST increased	1 (2%)	11 (22%)
Diarrhea	5 (10%)	10 (20%)
Oropharyngeal pain	3 (6%)	9 (18%)
Muscle spasm	3 (6%)	8 (16%)
Back pain	4 (8%)	7 (14%)
Decreased Appetite	2 (4%)	6 (12%)

¹AEs reported in >10% of omaveloxolone patients observed more frequently (>5% difference) in omaveloxolone compared to placebo.

²No new safety signals have been observed in the Extension study as of August 2021 data cut-off.

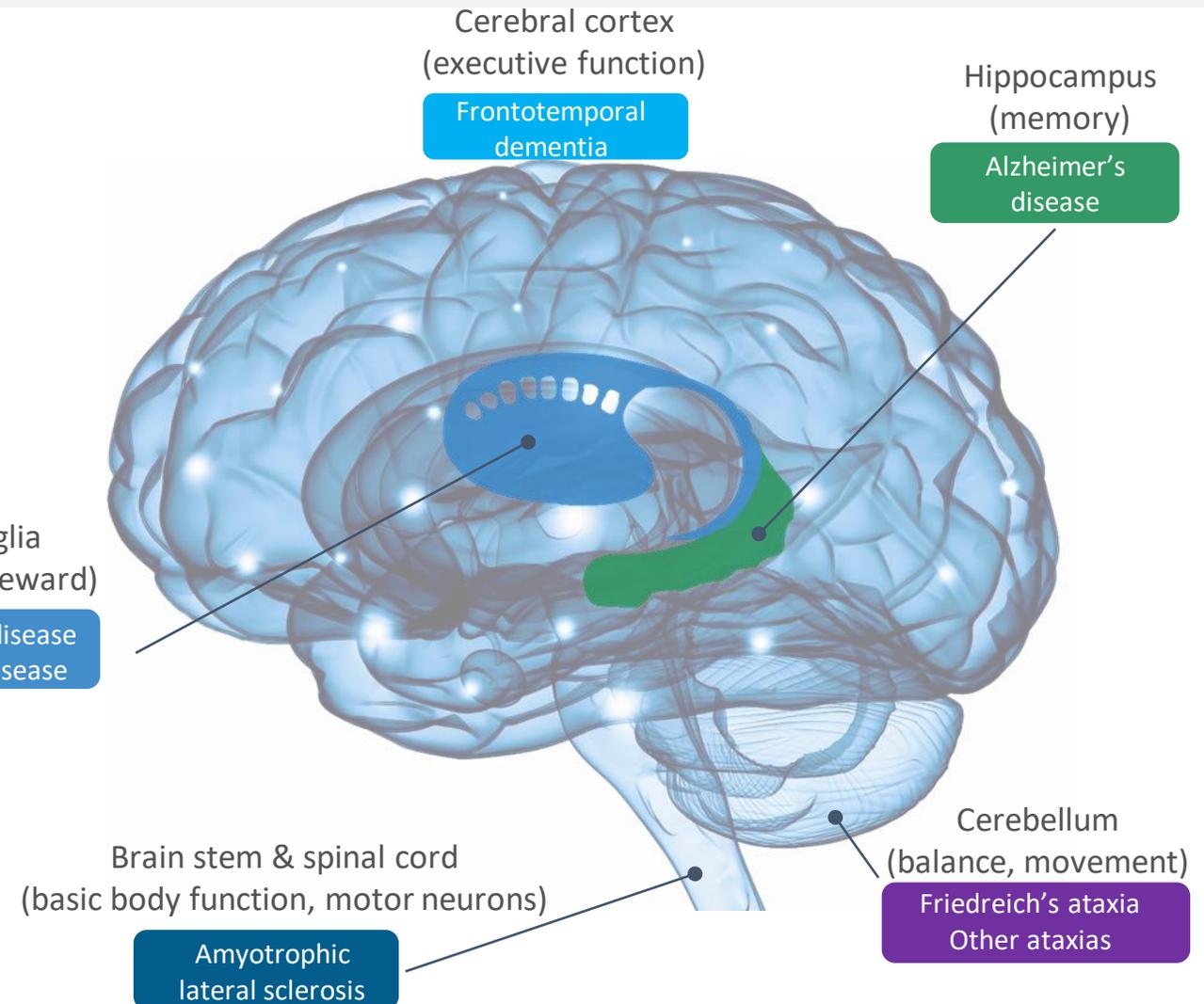
Omaveloxolone Pharmacology May Be **Applicable to a Broad Set of Neurological Diseases**

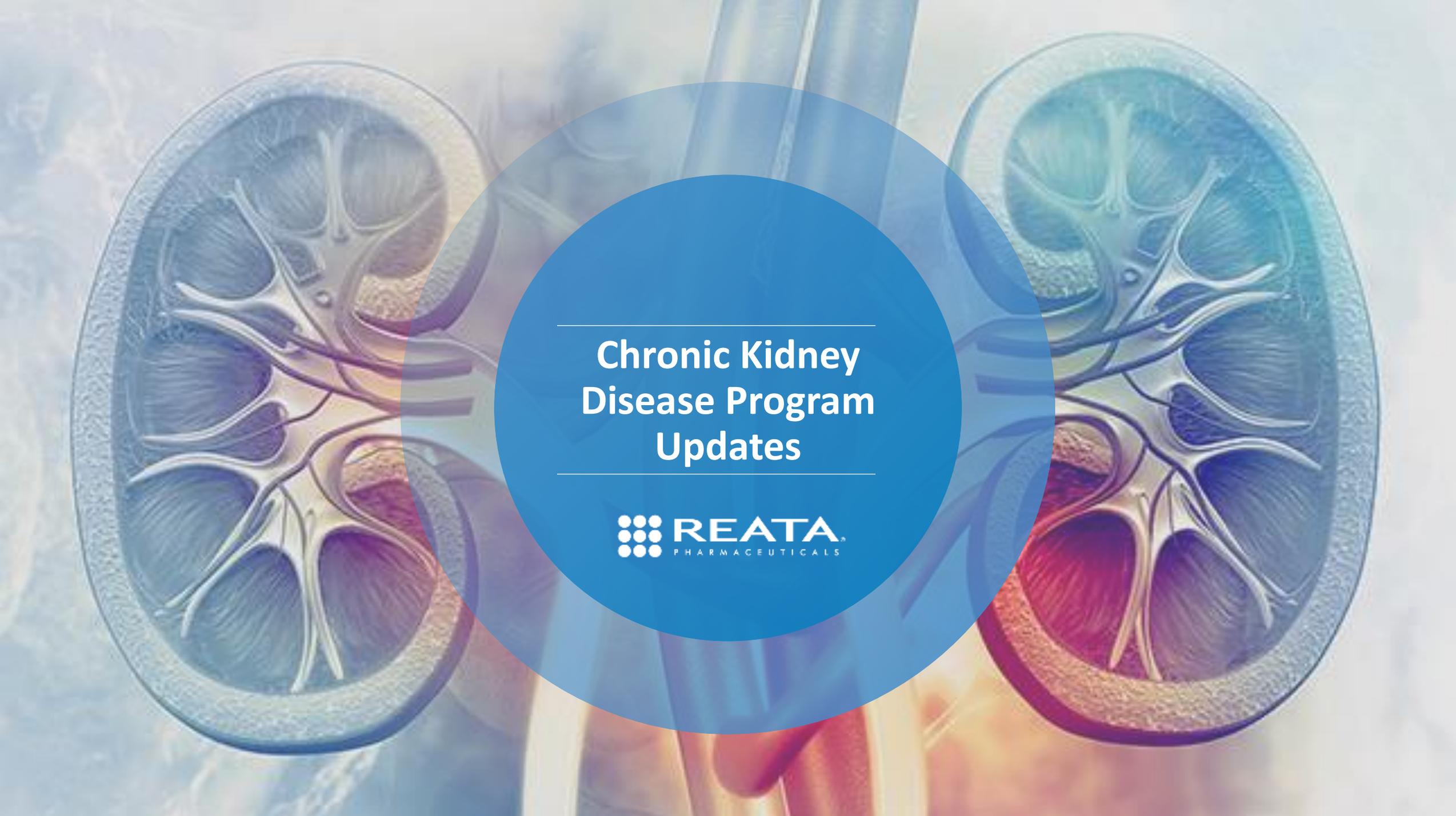
MOXIe results provide proof of concept for use of omaveloxolone in other neurological diseases

Mitochondrial dysfunction and neuroinflammation are common features of FA and other neurological diseases

Omaveloxolone and analogs may be applicable to the diseases listed below, and we have observed promising activity in preclinical models of many of these diseases

- Progressive supranuclear palsy
- Amyotrophic lateral sclerosis
- Parkinson's disease
- Frontotemporal dementia
- Huntington's disease
- Alzheimer's disease
- Epilepsy





Chronic Kidney Disease Program Updates

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AYAME Phase 3 Diabetic CKD Outcomes Trial in Japan

Kyowa Kirin, our strategic collaborator in Japan is sponsoring AYAME

Randomized, double-blind, placebo-controlled, registrational trial of bardoxolone in patients with diabetic CKD in Japan

- **Primary endpoint: time to onset of $\geq 30\%$ decline in eGFR or ESKD**
- Eligibility criteria
 - eGFR ≥ 15 and < 60 mL/min/1.73 m²
 - 20 to 79 years old
 - No history of heart failure or BNP > 200 pg/mL

More than 1,000 patients enrolled

Kyowa Kirin expects last patient visit in 2H 2022

Three years of data on all patients expected



FALCON Phase 3 Study of Bardoxolone in Patients with ADPKD

International, multi-center, randomized, double-blind, placebo-controlled Phase 3 trial of bardoxolone in patients with ADPKD

Protocol amendment filed with the FDA:

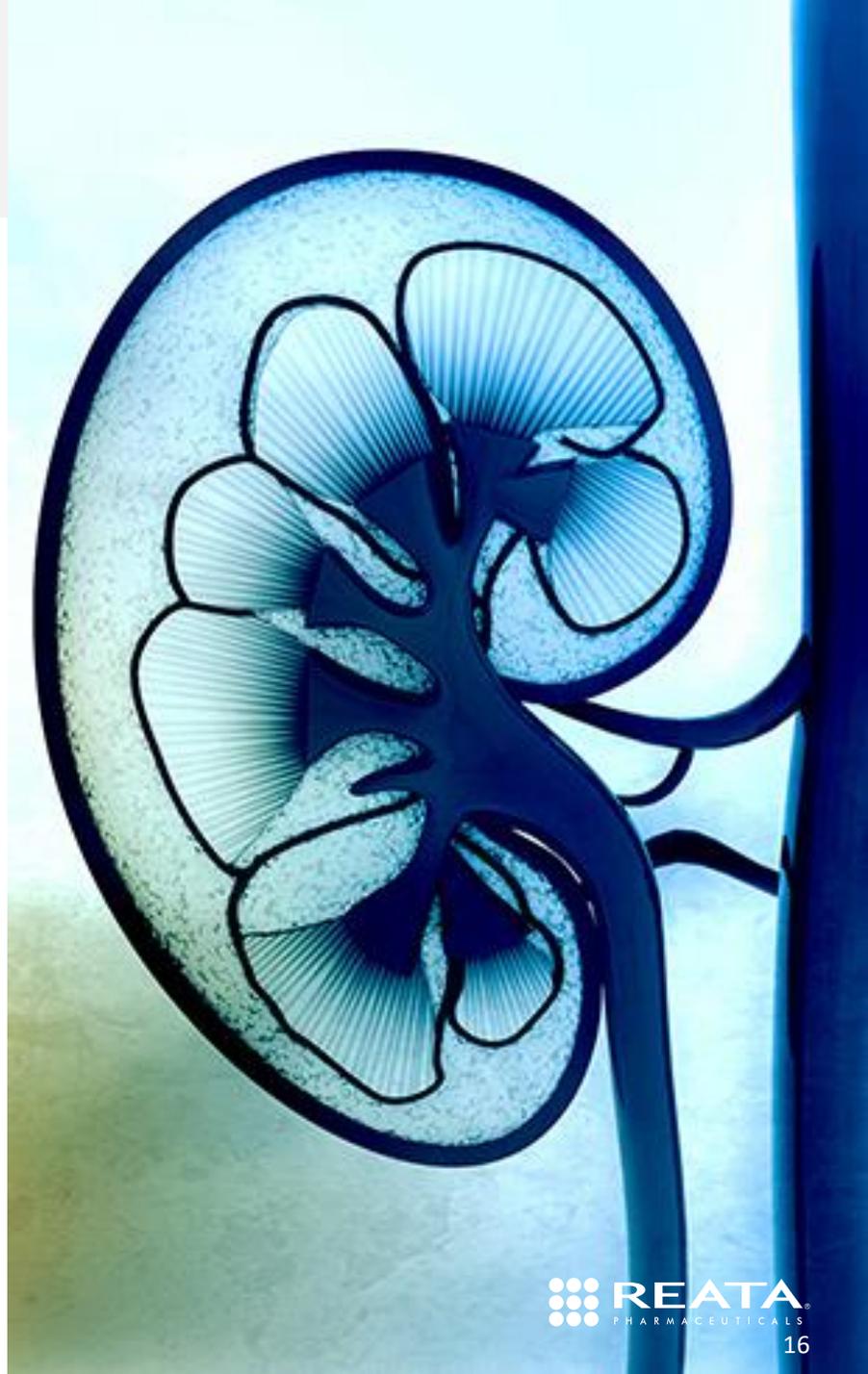
- **Endpoints**
 - **Primary:** off-treatment eGFR change from baseline at Week 108
 - No treatment discontinuation at Year 1
 - **Exploratory :** additional visit added to measure off-treatment eGFR at Week 112
- **Sample size**
 - Increased from 550 to 850 patients
- **Patient population**
 - Addition of adolescents ages 12 to 17 years (expanding age range - 12 to 70 years)
- **Others**
 - **Ambulatory blood pressure monitoring** sub-study added

>500 patients currently enrolled

Requested a Type A meeting with FDA to discuss overall ADPKD program including protocol amendment



ADPKD: Autosomal dominant polycystic kidney disease





RTA 901
HSP90 Modulator for
Diabetic Neuropathy

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RTA 901: Phase 2 Study in Patients with Diabetic Peripheral Neuropathic Pain (DPNP)

RTA 901 is a novel, small-molecule, orally bioavailable C-terminal Hsp90 inhibitor and Hsp70 inducer¹

DPNP is a serious complication of diabetes mellitus²⁻⁴

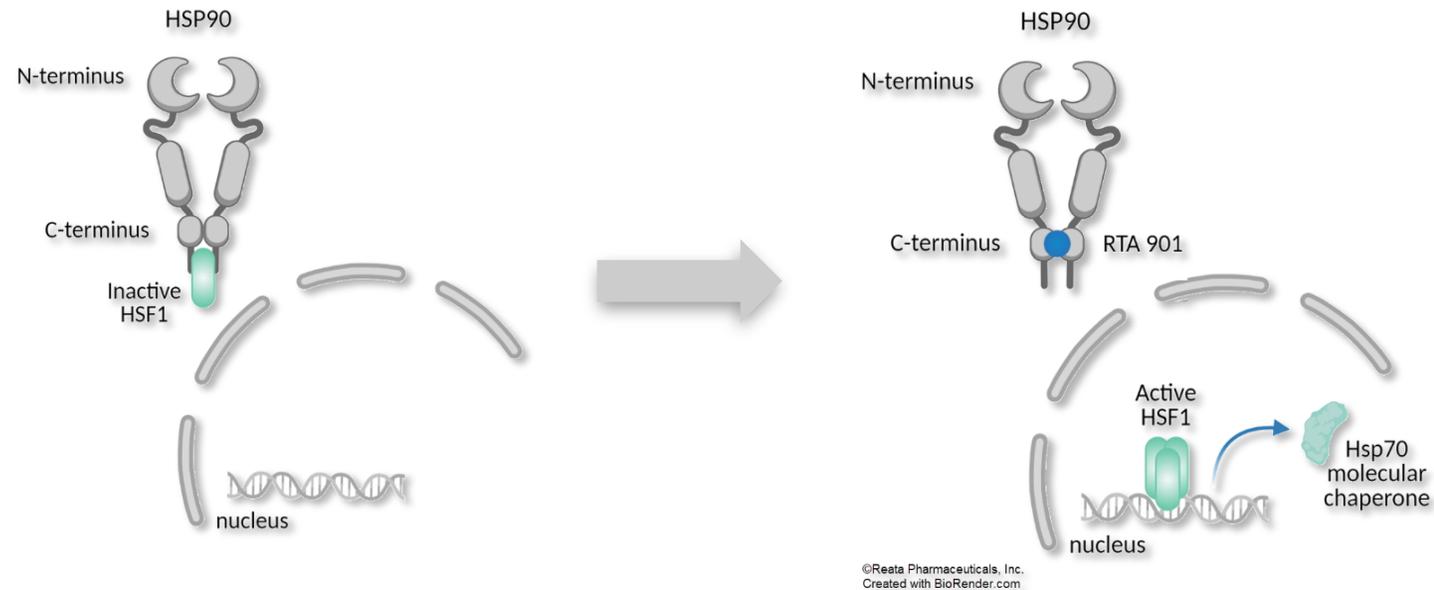
- Associated with substantial morbidity, including often debilitating peripheral neuropathic pain
- Approximately 4 million patients in the U.S. are affected with moderate or severe DPNP

Phase 1 study in healthy volunteers is complete

- Well tolerated with no safety signals, drug discontinuations, or SAEs
- Favorable PK profile

Additional Phase 1 clinical pharmacology studies expected to begin in the first half of 2022

Phase 2 randomized, placebo-controlled study in DPNP expected to begin enrolling in the second half of 2022



Financial & Operational Update



Significant U.S. Commercial Opportunity and Launch Readiness

Friedreich's Ataxia Launch Planning



Focused all commercial efforts to prepare for the launch of omaveloxolone



A small and efficient sales force will reach our target audience and customer targeting initiated



National accounts team deployed



Branded launch strategy development initiated

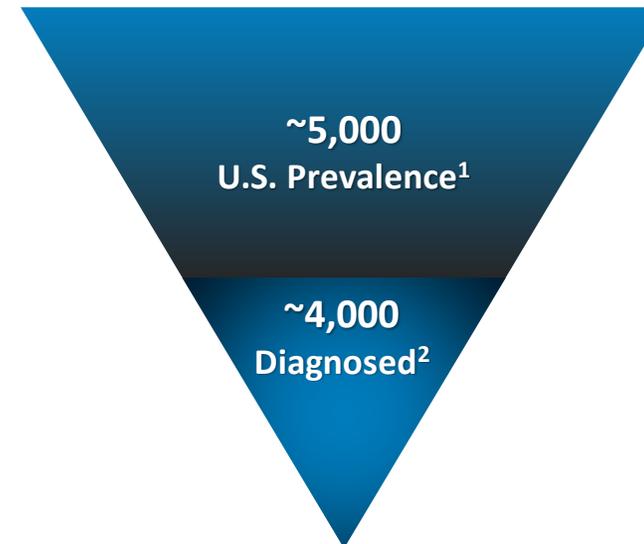


Friedreich's ataxia disease awareness continues

Friedreich's Ataxia Patients & Treatment Centers



U.S. Friedreich's Ataxia Patients



Centers for FA Patient Treatment

8
CCRN Centers³

22
U.S. Ataxia Centers³

225
MDA Clinics³

Key Neurology Practices

¹Estimated by Reata and the Friedreich's Ataxia Research Alliance (FARA) based on publicly available data and from Vankan P, J Neurochem 2013; ²2019 IQVIA claims data and projected diagnosed; ³Eight Collaborative Clinical Research Networks (CCRN) in Friedreich's Ataxia as disclosed by FARA, 22 ataxia centers as disclosed by the National Ataxia Foundation, and 225 Muscular Dystrophy Association (MDA) centers as disclosed by the Muscular Dystrophy Association

Financial Summary

	Three Months Ended December 31		Twelve Months Ended December 31	
	2021	2020	2021	2020
	(unaudited)			
Total Collaboration Revenue	\$ 0.9M	\$ 3.2M	\$ 11.5M	\$ 9.0M
Total GAAP Operating Expenses	\$ 72.5M	\$ 57.2M	\$ 256.2M	\$ 235.3M
Research and development	41.6M	37.5M	156.0M	159.1M
General and administrative	30.6M	19.4M	99.0M	75.1M
Depreciation	0.3M	0.3M	1.2M	1.1M
Non-GAAP Operating Expenses				
Non-GAAP Research and development ¹	35.5M	32.7M	132.4M	131.0M
Non-GAAP General and administrative ¹	21.4M	12.3M	65.8M	45.6M
GAAP Net Loss	\$ 85.4M	\$ 65.8M	\$ 297.4M	\$ 247.8M²
Non-GAAP Net Loss³	\$ 57.8M	\$ 43.5M	\$ 193.9M	\$ 158.3M²

Key Items at Dec 31, 2021

- Cash and Cash Equivalent \$590M
- Total Shares Outstanding 36.4M
 - 31.5M class A shares outstanding
 - 4.9M class B shares outstanding

Updated Cash Guidance:

- Based on our operational plans, we are extending cash guidance from mid-2024 to the end of 2024

¹Excludes stock-based compensation expenses; ²GAAP and non-GAAP net loss in the first quarter of 2020 includes the recognition of \$22.2 million tax benefits; ³Excludes various adjustments, including stock-based compensation expenses, non-cash interest expense from liability related to sale of future royalties, loss on extinguishment of debt, gain on lease termination; see the next slide for a reconciliation between GAAP and non-GAAP measures. GAAP: Generally Accepted Accounting Principles; M: Million

Reconciliation of GAAP to Non-GAAP Financial Measures

	Three Months Ended December 31		Twelve Months Ended December 31	
	2021	2020	2021	2020
Reconciliation of GAAP to Non-GAAP Research and development:				
GAAP Research and development	\$ 41,616	\$ 37,461	\$ 155,993	\$ 159,080
Less: Stock-based compensation expense	(6,091)	(4,792)	(23,566)	(28,114)
Non-GAAP Research and development	\$ 35,525	\$ 32,669	\$ 132,427	\$ 130,966
Reconciliation of GAAP to Non-GAAP General and administrative:				
GAAP General and administrative	\$ 30,562	\$ 19,427	\$ 99,002	\$ 75,128
Less: Stock-based compensation expense	(9,135)	(7,158)	(33,240)	(29,519)
Non-GAAP General and administrative	\$ 21,427	\$ 12,269	\$ 65,762	\$ 45,609
Reconciliation of GAAP to Non-GAAP Operating expenses:				
GAAP Operating expenses	\$ 72,503	\$ 57,173	\$ 256,198	\$ 235,344
Less: Stock-based compensation expense	(15,226)	(11,950)	(56,806)	(57,633)
Non-GAAP Operating expenses	\$ 57,277	\$ 45,223	\$ 199,392	\$ 177,711
Reconciliation of GAAP to Non-GAAP Net loss:				
GAAP Net loss	\$ (85,385)	\$ (65,776)	\$ (297,386)	\$ (247,752)
Add: Stock-based compensation expense	15,226	11,950	56,806	57,633
Add: Non-cash interest expense from liability related to sale of future royalties	12,376	10,807	46,688	21,884
Add: Loss on extinguishment of debt	-	-	-	11,183
Less: Gain on lease termination	-	(470)	-	(1,286)
Non-GAAP Net loss	\$ (57,783)	\$ (43,489)	\$ (193,892)	\$ (158,338)
Reconciliation of GAAP to Non-GAAP Net loss per common share-basic and diluted:				
GAAP Net loss per common share-basic and diluted	\$ (2.35)	\$ (1.90)	\$ (8.19)	\$ (7.35)
Add: Stock-based compensation expense	0.42	0.35	1.56	1.71
Add: Non-cash interest expense from liability related to sale of future royalties	0.34	0.31	1.29	0.65
Add: Loss on extinguishment of debt	-	-	-	0.33
Less: Gain on lease termination	-	(0.01)	-	(0.04)
Non-GAAP Net loss per common share-basic and diluted	\$ (1.59)	\$ (1.25)	\$ (5.34)	\$ (4.70)



Concluding Remarks



Reata at a Glance



Neurology Pipeline

- Initiated rolling submission of NDA for omaveloxolone in FA
- Plan to complete submission by the end of Q1 2022
- Plan to initiate Phase 2 study of RTA 901 in DPNP in second half of 2022



CKD Pipeline

- Received CRL and will work with FDA to confirm next steps on Alport program
- FALCON protocol amendment submitted to FDA and requested Type A meeting
- >500 patients enrolled in FALCON



Global Opportunity

- Strong cash position
- Worldwide commercial rights to all pipeline assets¹
- Robust IP protection for omaveloxolone, RTA 901, and bardoxolone

¹Ex-Asia for bardoxolone IP: Intellectual Property



Thank you



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