



# **Annual General Meeting of Shareholders**

May 10<sup>th</sup> 2017

# Forward looking statements

This presentation contains forward-looking statements that involve substantial risks and uncertainties. All statements, other than statements of historical facts, contained in this presentation, including but not limited to, statements regarding our strategy, future operations, future pre-clinical and clinical trial plans and related timing of trials and results, research and development, future financial position, future revenues, projected costs, prospects, therapeutic potential of our products, plans and objectives of management, are forward-looking statements. The words “aim,” “anticipate,” “believe,” “estimate,” “expect,” “intend,” “may,” “plan,” “predict,” “project,” “target,” “potential,” “will,” “would,” “could,” “should,” “continue,” and similar expressions are intended to identify forward-looking statements, although not all forward-looking statements contain these identifying words.

Forward-looking statements represent our management’s beliefs and assumptions only as of the date of this presentation. We may not actually achieve the plans, intentions or expectations

disclosed in our forward-looking statements, and you should not place undue reliance on our forward-looking statements. Actual results or events could differ materially from the plans, intentions and expectations disclosed in the forward-looking statements we make. The forward-looking statements contained in this presentation reflect our current views with respect to future events, and we assume no obligation to update any forward-looking statements except as required by applicable law. These forward-looking statements are subject to a number of risks, uncertainties and assumptions, including those that may be described in greater detail in the annual report filed on Form 20-F for the year ended December 31, 2016 that we have filed with the U.S. Securities and Exchange Commission (the “SEC”) and any subsequent filings we have made with the SEC. We have included important factors in the cautionary statements included in that annual report, particularly in the Risk Factors section, and subsequent filings with the SEC that we believe could cause actual results or events to differ materially from the forward-looking statements that we make.

# 1. Opening of the AGM

Antoine Papiernik, Vice Chairman of the Supervisory Board



# Agenda

1. Opening of the AGM
2. Report of the Management Board
3. Disclosure of remuneration in the annual accounts for the financial year 2016

## **Voting items:**

- 4. Adoption of the annual accounts for the financial year 2016**
- 5. Release from liability of the members of the Management Board**
- 6. Release from liability of the members of the Supervisory Board**
- 7. Reappointment of Supervisory Board member Antoine Papiernik**
- 8. Appointment Deloitte Accountants B.V.**
- 9. Delegation to the Management Board of the authority (i) to issue ordinary shares, (ii) to grant rights to subscribe for such shares and (iii) to limit and exclude pre-emption rights**
- 10. Authorization of the Management Board to acquire ordinary shares**
11. Questions
12. Closing of the AGM

## **2. Discussion Item**

# **Report of the Management Board for the financial year 2016**

Daniel de Boer, Chief Executive Officer

# 2016 overview

*From idea to proof-of-concept in CF patients*

- Translated an idea for CF into clinical PoC in patients
- Expanded from 1 to 3 development programs
  - QR-010 for cystic fibrosis
  - QR-110 for Leber's congenital amaurosis type 10
  - QR-313 for dystrophic epidermolysis bullosa
- Discovery pipeline to fuel the future



# Our mission and strategy



## Treat Genetic Diseases

- ProQR was founded to find a treatment for CF
- Less than 10% of genetic diseases have a treatment
- Creating treatments for severe rare diseases where we can have a big impact



## Build strong foundation

- Experienced team with proven track record
- Top-notch science and collaborators
- Broad IP estate



## RNA therapeutics

- Elegant & highly targeted approach
- Emergent RNA field with 5 approved products
- Established modality through >20 years of experience (delivery, safety, manufacturing, etc)



## Created valuable pipeline

- Diversified product pipeline
- Focused on high unmet medical need and accelerated development pathways



## Build a sustainable business

- Best-in-class products
- Achievable commercial strategy in rare diseases
- Drive shareholder value

# Strong team with proven track record

## Management team



Daniel de Boer  
*Chief Executive Officer*



Noreen Henig  
*Chief Medical Officer*



Gerard Platenburg  
*Chief Innovation Officer*



Smital Shah  
*Chief Financial Officer*



René Beukema  
*Chief Corp. Development Officer & General Counsel*



Robert Cornelisse  
*Chief People & Organization*



David Rodman  
*Chief Development Strategy Officer*



## Supervisory board



Dinko Valerio  
*Chairman*



Henri Termeer



James Shannon



Alison Lawton



Paul Baart



Antoine Papiernik





# Key collaborators

*In academia and patient organizations*



MASSACHUSETTS  
GENERAL HOSPITAL

Radboud  
Universiteit  
Nijmegen



ROSALIND FRANKLIN  
UNIVERSITY  
of MEDICINE AND SCIENCE



THE UNIVERSITY  
of NORTH CAROLINA  
at CHAPEL HILL



Stanford  
University



Leids Universitair  
Medisch Centrum



University of  
BRISTOL

FOUNDATION  
FIGHTING  
BLINDNESS

debra  
because the cost of doing nothing is too great

CYSTIC FIBROSIS  
FOUNDATION

# Therapeutic Strategy



## Patient centric

Best-in-class high impact products for patients in need



## Well understood causality

Single gene defect leading to disease manifestation



## Local delivery

Feasible delivery route to target organ



## Genetic rare diseases

Limited treatment options, viable commercial strategy



## RNA therapy

Highly specific approach for a wide range of mutations



## Established modality

Matured RNA field, proven best practices

# Pipeline in key therapeutic areas



**Cystic  
fibrosis**



**Inherited  
blindness**



**Debilitating skin  
disorders**







# Cystic fibrosis



**High unmet  
medical need**



**Lung &  
other organs**



**Limited life  
expectancy of  
27 years**



**Most common  
mutation affects  
~65,000 patients**

# QR-010 for cystic fibrosis

*Differentiating product profile*



## **INHALED DRUG**

for lung delivery and systemic uptake



## **SINGLE AGENT for F508del**

to treat underlying cause of disease



## **CONVENIENT AT HOME DOSING**

3 times a week or less in under 15 minutes



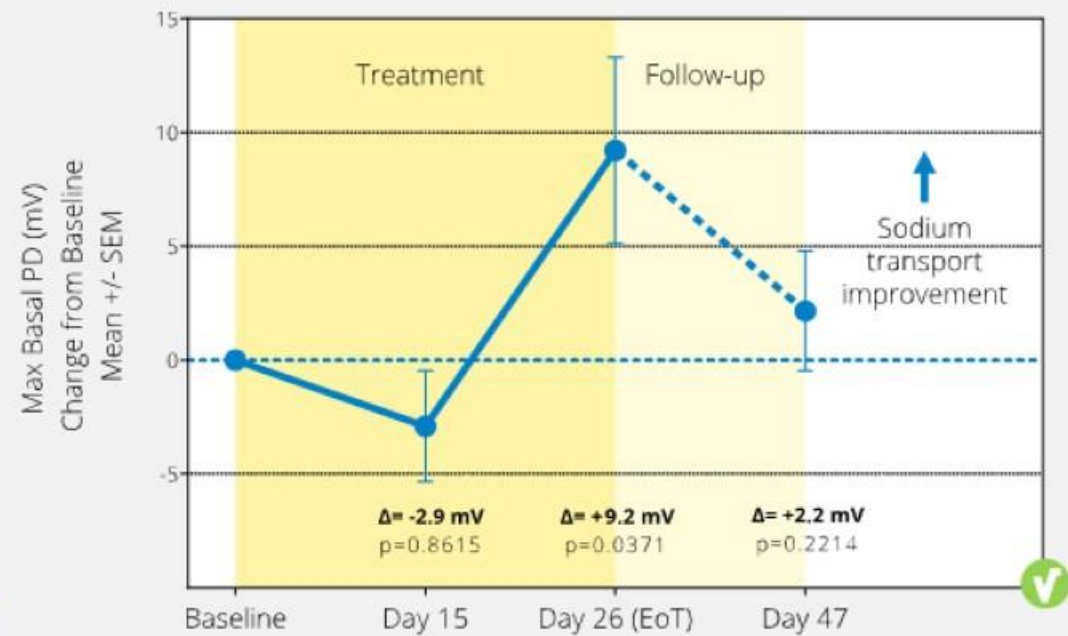
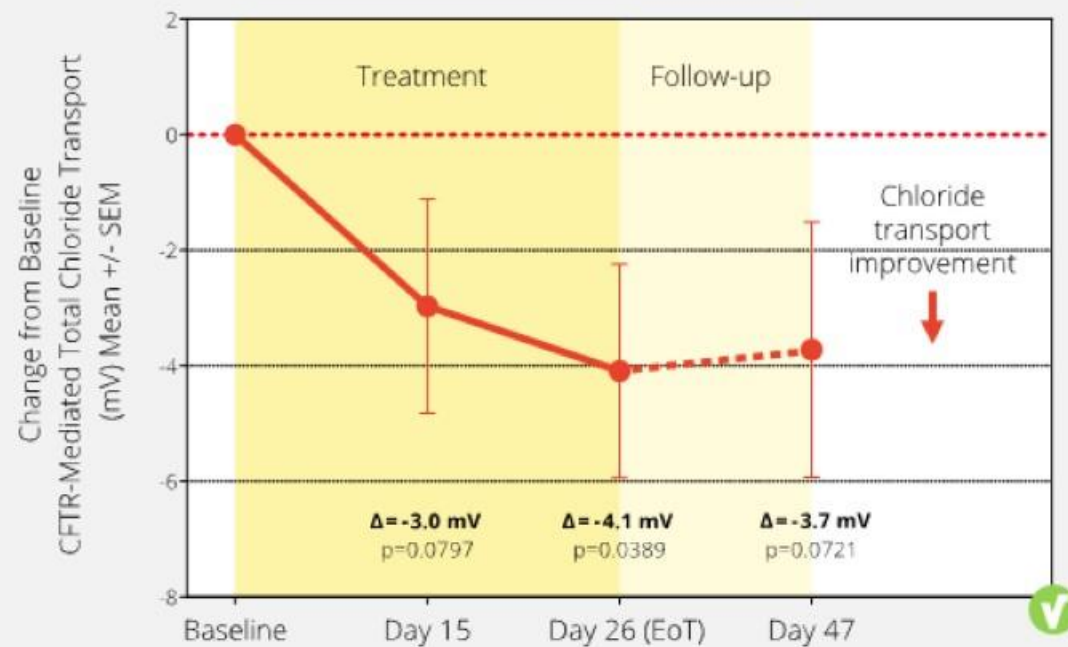
## **AIMS TO STOP PROGRESSION OF DISEASE**

or prevent disease and improve quality of life



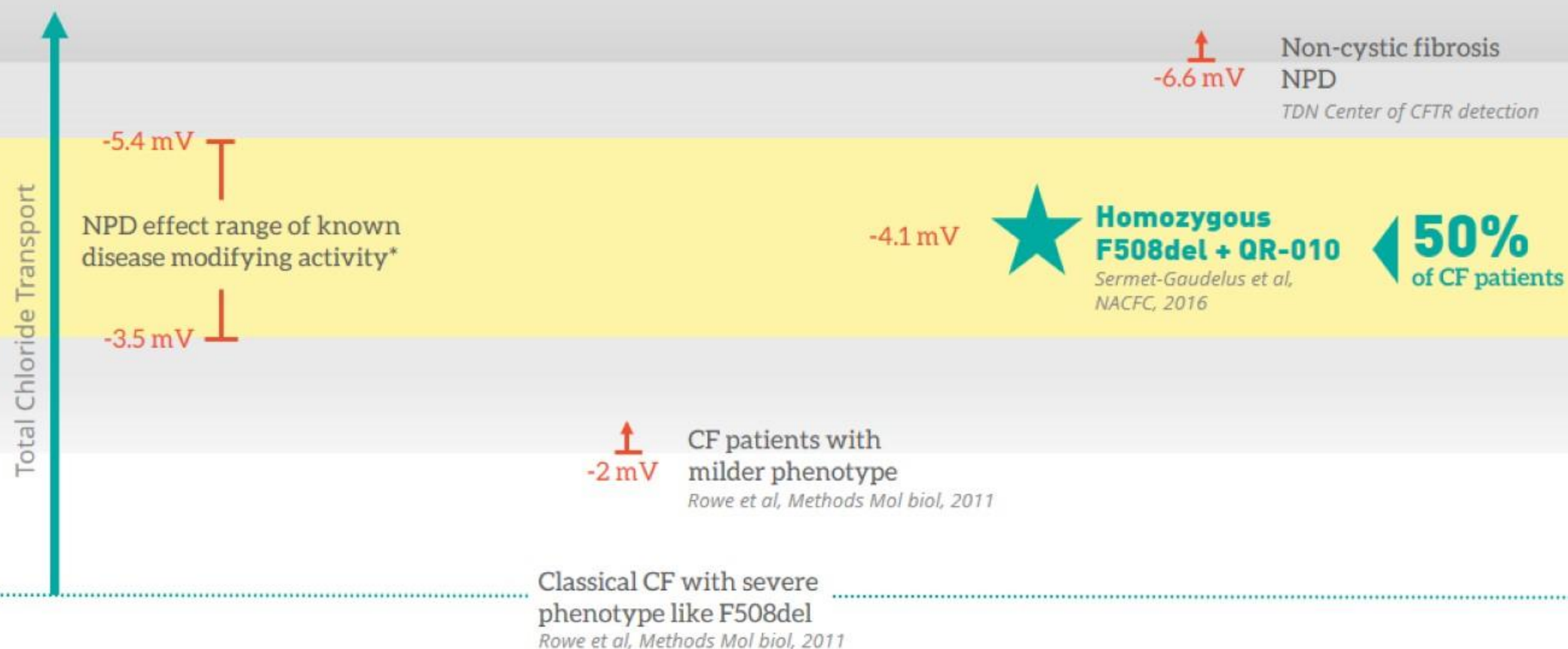
Investigational product. Not available for commercial distribution in the U.S.

# QR-010 improves CFTR activity in CF patients





# QR-010: CFTR levels that are expected to be disease modifying



Interpretations are adapted from publications

\* Based on responses in ivacaftor studies in G551D

# Bringing QR-010 to CF patients

## Safety study PQ-010-001



### Single ascending dose cohorts

- All 4 cohorts completed
- QR-010 was well tolerated without safety concerns

### Global Phase 1b trial

- 64 homozygous F508del patients with good lung function (high FEV<sub>1</sub>)



### Multiple ascending dose cohorts

- 2 cohorts completed
- Study completion in mid-2017

- 27 sites in Europe and North America
- Leading CF investigators and centers participating

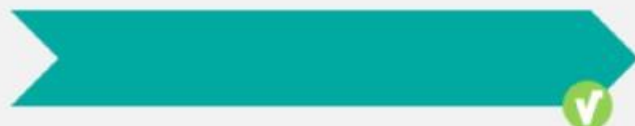
## Efficacy PQ-010-003



### Safety, PK and efficacy study to begin enrolment in 2018

- 12-week, placebo controlled in more severe CF patients (lower FEV<sub>1</sub>)
- Global study with sites in North America, EU and other countries
- Efficacy measurements under consideration: FEV<sub>1</sub>, LCI, Sweat, CFQ-R, Weight, Imaging, Biomarkers

## Drug activity study PQ-010-002



### Nasal potential difference biomarker study

- QR-010 restored CFTR function in homozygous F508del CF patients

# QR-010

## “Stop the progression of cystic fibrosis”

- ✓ Drug activity as monotherapy
- ✓ Established proof of concept in CF patients
- ✓ Rapid progression from pre-clinical to clinical stage (3 years)
- ✓ Fast track designation

### Next steps

- Completion of Phase 1b study in mid 2017
- Phase 2 study initiation in 2018 driving QR-010 to CF patients







# Inherited blindness

*Restoring vision or stop the progression of inherited blindness.*

**>300**

**Genetic eye  
diseases without  
treatment**



**Pipeline in  
several inherited  
blindness**



**Established  
modality in eye**

## Leber's congenital amaurosis

*QR-110: First product in pipeline targets LCA 10 (p.Cys998X)*



**Lose sight in first  
years of life**



**No therapy  
available**



**LCA 10 affects  
~2.000 patients in  
western world**

# QR-110 for LCA 10



**AIMS TO RESTORE VISION/PREVENT VISION LOSS IN PATIENTS SUFFERING FROM LCA 10**



**LOCALLY ADMINISTERED IN THE EYE**

Routine intravitreal procedure

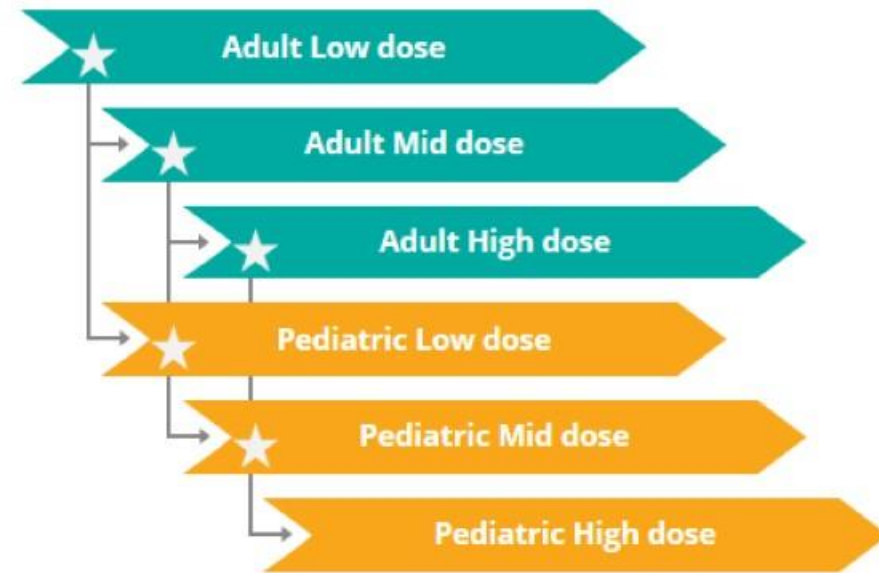


**INFREQUENT DOSING**

4 times a year or less



# QR-110: Phase 1/2 safety & efficacy trial in LCA 10 patients



2 patients per cohort,  
open-label

☆ DSMC review

- 6 adults and 6 children (≥6yrs) homozygous or compound heterozygous for p.Cys998X mutation
- Intravitreal injections in one eye, other eye and subject's own baseline serve as controls
- Dosing scheme: every 3 months, maximum 4 doses
- Participating sites: 3 major sites in EU and US
- Primary objectives: Safety, tolerability
- Secondary objectives: Pharmacokinetics, efficacy (visual acuity, FST, OCT, PLR, mobility course)
- Inclusion criteria: detectable ONL, ERG consistent with LCA, possibility for retinal imaging, some visual acuity
- **Upcoming updates:** 1<sup>st</sup> patient dosed, enrollment complete, top-line results (expected in 2018)





# Debilitating skin diseases

*dystrophic epidermolysis bullosa*



**Limited life expectancy**



**Building a pipeline to help all DEB patients**



**Skin & other organs**



**Blistering from birth**



**First product targets exon 73**



**Most common mutation affects ~2,000 patients**

# QR-313 for dystrophic epidermolysis bullosa



**AIMS TO HEAL WOUNDS, RESTORE SKIN  
AND IMPROVE QUALITY OF LIFE**



## **TOPICALLY APPLIED**

Commonly used hydrogel, containing  
QR-313 RNA therapy



## **CONVENIENT APPLICATION AT HOME**

Maximum frequency every other day



# First-in-human clinical trail in DEB patients

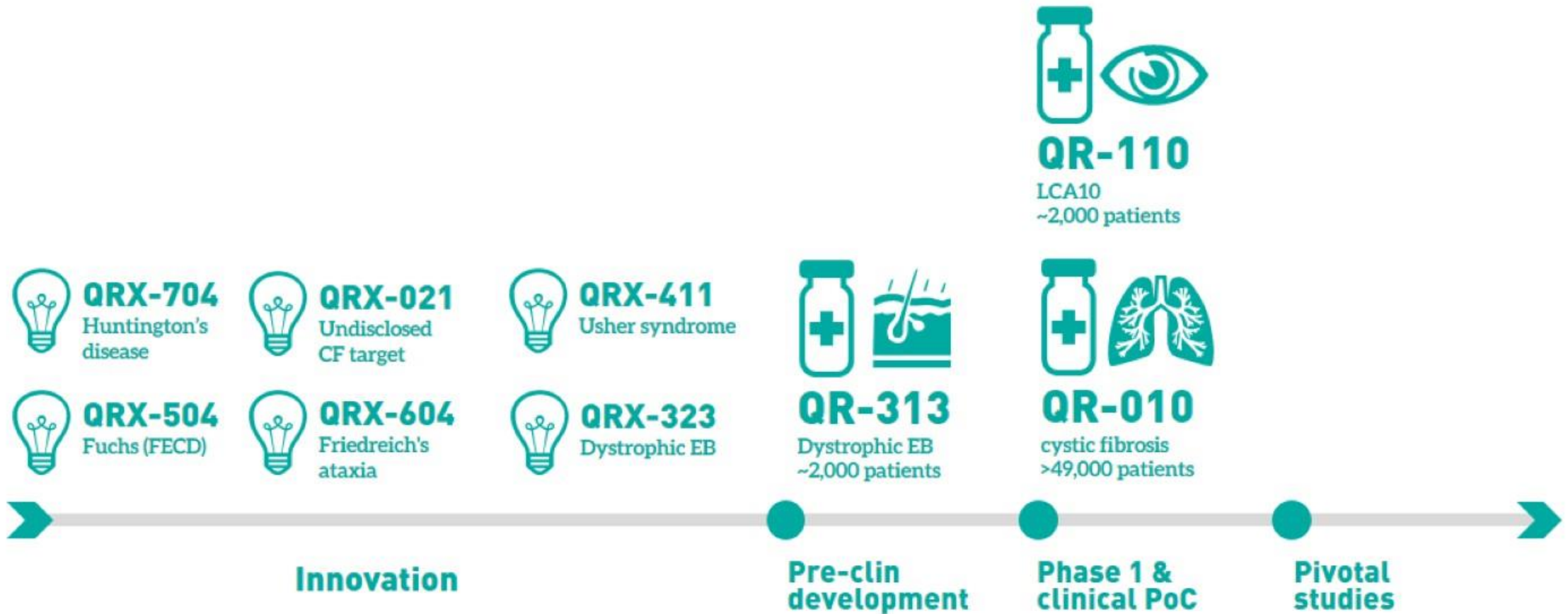
## QR-313 leading the pipeline for DEB

- Up to 3 month study in DEB patients
- Up to 10 patients with exon 73 mutation
- Clinical endpoints: Safety and tolerability, exploratory efficacy and biomarkers
- Up to 5 sites in 3 countries
- **Expected timing:** Dose 1st patients in 2018, top-line data in 2018
- Translating QR-313 into pipeline compounds for other DEB mutations





# And beyond



# Rapidly advancing our products to patients



## QR-010 for cystic fibrosis

- Top-line results of 64 patient Phase 1b trial in mid-2017
- Start phase 2 program in 2018



## QR-110 for Inherited blindness LCA 10

- Starting clinical trial in adult and pediatric patients in H1 2017
- Top-line results of clinical trial in 2018



## QR-313 for debilitating skin disease DEB

- Conducting IND enabling studies in 2017
- Start and readout clinical trial in DEB patients in 2018







# Financial highlights

Smital Shah, Chief Financial Officer

# Balance Sheet – Annual Changes

	DECEMBER 31, 2016	DECEMBER 31, 2016
	€1000	€1000
<b>ASSETS</b>		
<b>Non-current assets</b>	<b>3,528</b>	<b>2,340</b>
Intangible assets	90	141
Property, plant and equipment	3,438	2,199
<b>Current assets</b>	<b>62,015</b>	<b>97,769</b>
Social securities and other taxes	395	956
Prepayments and other receivables	2,420	1,948
Cash and cash equivalents	59,200	94,865
<b>TOTAL ASSETS</b>	<b>65,543</b>	<b>100,109</b>
<b>EQUITY &amp; LIABILITIES</b>		
<b>Shareholders' equity</b>	<b>53,136</b>	<b>89,799</b>
<b>Non-current liabilities</b>	<b>5,697</b>	<b>4,824</b>
Finance lease liabilities	--	--
Borrowings	5,697	4,824
<b>Current liabilities</b>	<b>6,710</b>	<b>5,486</b>
Finance lease liabilities	--	15
Trade payables	328	885
Social securities and other taxes	312	235
Pension premiums	13	16
Deferred income	--	144
Other current liabilities	6,057	4,191
<b>TOTAL EQUITY AND LIABILITIES</b>	<b>65,543</b>	<b>100,109</b>

# P&L – Annual Changes

	2016	2015
	€ 1,000	€ 1,000
<b>Other income</b>	<b>1,828</b>	<b>3,235</b>
Research and development costs	(31,923)	(23,401)
General and administrative costs	(9,478)	(6,837)
<b>Total operating costs</b>	<b>(41,401)</b>	<b>(30,238)</b>
<b>Operating result</b>	<b>(39,573)</b>	<b>(27,003)</b>
Financial income and expense	470	6,171
<b>Result before corporate income taxes</b>	<b>(39,103)</b>	<b>(20,832)</b>
Corporate income taxes	--	--
<b>Result for the year</b>	<b>(39,103)</b>	<b>(20,832)</b>
<b>Other comprehensive income</b>	<b>(16)</b>	<b>1</b>
<b>Total comprehensive income for the year (attributable to equity holders of the Company)</b>	<b>(39,119)</b>	<b>(20,831)</b>
<b>Earnings per share attributable to the equity holders of the Company (expressed in Euro per share)</b>		
Basic earnings per share <sup>1</sup>	(1.68)	(0.89)
Diluted earnings per share <sup>1</sup>	(1.68)	(0.89)



# Cash Flow – Annual Changes

	2016	2015
	€ 1,000	€ 1,000
Result for the year	(39,119)	(20,831)
Adjustments for:		
— Depreciation	1,245	480
— Share-based compensation	2,454	1,212
— Financial income and expense	(470)	(6,171)
Changes in working capital	1,433	637
Cash used in operations	(34,457)	(24,673)
Corporate income tax paid	--	--
Interest received/(paid)	236	441
<b>Net cash used in operating activities</b>	<b>(34,221)</b>	<b>(24,232)</b>
Purchases of intangible assets	--	(28)
Purchases of property, plant and equipment	(2,539)	(1,296)
<b>Net cash used in investing activities</b>	<b>(2,539)</b>	<b>(1,324)</b>
Proceeds from exercise of share options	2	14
Proceeds from borrowings	370	1,640
Redemption of financial lease	(15)	(34)
<b>Net cash generated by financing activities</b>	<b>357</b>	<b>1,620</b>
<b>Net increase/(decrease) in cash and cash equivalents</b>	<b>(36,403)</b>	<b>(23,936)</b>
Currency effect cash and cash equivalents	738	6,065
Cash and cash equivalents at the beginning of the year	94,865	112,736
<b>Cash and cash equivalents at the end of the year</b>	<b>59,200</b>	<b>94,865</b>

### **3. Discussion Item**

## **Disclosure of remuneration in the annual accounts for the financial year 2016**

# Disclosure of remuneration in the annual accounts for 2016 (i)

## Remuneration of Management Board in 2014

Name	Short term employee benefits	Post Employment benefit	Share-based payment	Total
	(€1000)	(€1000)	(€1000)	(€1000)
Daniel de Boer	429	7	391	827
Rene Beukema	346	13	165	524



# Disclosure of remuneration in the annual accounts for 2016 (ii)

## Remuneration of Supervisory Board in 2016

Name	Short term employee benefits	Post Employment benefit	Share-based payment	Total
	(€1000)	(€1000)	(€1000)	(€1000)
Dinko Valerio	36	--	52	88
Henri Termeer	31	--	51	82
Antione Papiernik	78	--	--	78
Alison Lawton	31	--	74	105
Paul Baart	82	--	--	82
James Shannon	29	--	27	56

## **4. Voting Item**

# **Adoption of the annual accounts for the financial year 2016**

## 4. Voting Item

# Adoption of the annual accounts for the financial year 2016

Votes For	Votes Against	Votes Abstained
14,438,891	3,450	10,032



## 5. Voting Item

**Release from liability of the members of the Management Board with respect to the performance of their management during the financial year 2016**

## 5. Voting Item

**Release from liability of the members of the Management Board with respect to the performance of their management during the financial year 2016**

Votes For	Votes Against	Votes Abstained
13,736,328	206,372	509,673

## **6. Voting Item**

**Release from liability of the members of the Supervisory Board with respect to the performance of their supervision during the financial year 2016**



## 6. Voting Item

**Release from liability of the members of the Supervisory Board with respect to the performance of their supervision during the financial year 2016**

Votes For	Votes Against	Votes Abstained
13,736,328	206,372	509,673

## **7. Voting Item**

# **Reappointment of Supervisory Board member Antoine Papiernik**

# Composition Supervisory Board

Name	Term – until
Henri Termeer	2020
Dinko Valerio	2020
Antoine Papiernik	2017
Alison Lawton	2018
Paul Baart	2019
James Shannon	2020



## 7. Voting Item

# Reappointment of Supervisory Board member Antoine Papiernik

Votes For	Votes Against	Votes Abstained
14,442,614	6,550	3,209

## **8. Voting Item**

**Appointment Deloitte Accountants  
B.V. as the company's external  
auditor for the financial year 2018**

## 8. Voting Item

# Appointment Deloitte Accountants B.V. as the company's external auditor for the financial year 2018

Votes For	Votes Against	Votes Abstained
14,445,523	6,550	300



## 9. Voting Item

**Delegation to the Management Board of the authority (i) to issue ordinary shares, (ii) to grant rights to subscribe for such shares and (iii) to limit and exclude pre-emption rights**

# Authorization to issue shares

## **Proposed authorization of Management Board:**

- a) To issue ordinary shares for general purposes and/or for mergers, demergers, acquisitions and other strategic transactions and alliances (or a combination thereof) up to 30% of the Company's issued share capital, plus for issuance under stock option plans up to 15% of the Company's issued share capital (minus any treasury shares),
  - b) To grant rights to subscribe for ordinary shares as described under (a)
  - c) To limit or exclude the pre-emptive rights of holders of ordinary shares
- Valid for a period of 5 years from today
  - Includes the authority to determine the price and further terms and conditions of any such share issuance or grant
  - Subject to approval of the Supervisory Board



## 9. Voting Item

**Delegation to the Management Board of the authority (i) to issue ordinary shares, (ii) to grant rights to subscribe for such shares and (iii) to limit and exclude pre-emption rights**

Votes For	Votes Against	Votes Abstained
12,610,620	979,951	861,802



## **10. Voting Item**

**Authorization of the Management Board to acquire ordinary shares in the capital of the Company**

# Authorization to acquire shares

## **Proposed authorization of Management Board:**

- To acquire up to 10% of issued shares
- In case of material reorganization of capital structure an additional 10%
- Authorization lasts for 18 months after AGM

## 10. Voting Item

### Authorization of the Management Board to acquire ordinary shares in the capital of the Company

Votes For	Votes Against	Votes Abstained
13,139,258	1,312,715	400

# 11. Questions & Answers



# **12. Closing of the AGM**



**IT'S IN  
OUR RNA**