

2019 OIS@ASCRS May 2nd , 2019

Innovative Solutions
to Help Maintain Vision
and Improve Ocular Health

Euronext Paris: COX

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Risk factors which are likely to have a material effect on Nicox SA's business are presented in the 4th chapter of the "*Document de référence, rapport financier annuel et rapport de gestion 2018*" filed with the French *Autorité des Marchés Financiers* (AMF) on March 6, 2019 under number D.19-0117 available on Nicox SA's website (www.nicox.com).

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Nicox: Why We Are a Unique Ophthalmology R&D Company

RESEARCH

*Two new innovative classes
of NO-donating drug candidates
for glaucoma*

DEVELOPMENT

*NCX 470 - Potential “best in class”
mono-compound for glaucoma
NCX 4251 – Potential “first in class”
treatment for blepharitis*

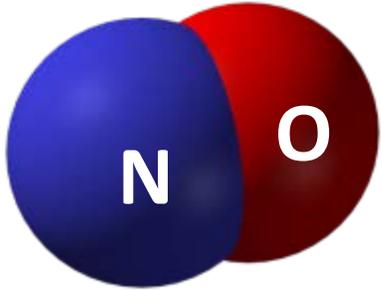
MARKETING

*International expansion
of two FDA approved products,
VYZULTA® and ZERVIATE™*

Broad Pipeline of Ophthalmic Therapeutics

Products and product candidates / Indications	Rights	Stages of Development						
		Research	Preclinical	Phase 1	Phase 2	Phase 3	NDA	Marketed
NO-Donating Product Candidates Targeting Glaucoma and Other Indications								
NCX 470 Second generation NO-donating PGA Glaucoma	  ¹	[Progress bar from Research to Phase 2]						
Future generation NO-donors Glaucoma		[Progress bar from Research to Preclinical] NO-donating sGC stimulators						
		[Progress bar from Research to Preclinical] NO-donating PDE5 inhibitors						
Novel Formulation Targeting Blepharitis								
NCX 4251 fluticasone propionate Blepharitis		[Progress bar from Research to Phase 1]						
Out-Licensed Commercial Products and Product Candidate								
VYZULTA® Glaucoma	 BAUSCH + LOMB ²	[Progress bar from Research to Marketed]						
ZERVIAE™ Allergic conjunctivitis	 ³  ⁴ 	[Progress bar from Research to Phase 3]						
NCX 4280 Morning ocular congestion		[Progress bar from Research to Phase 2]						

The Central Role of Nitric Oxide in IOP Homeostasis



Nobel Prize 1998

RF Furchgott

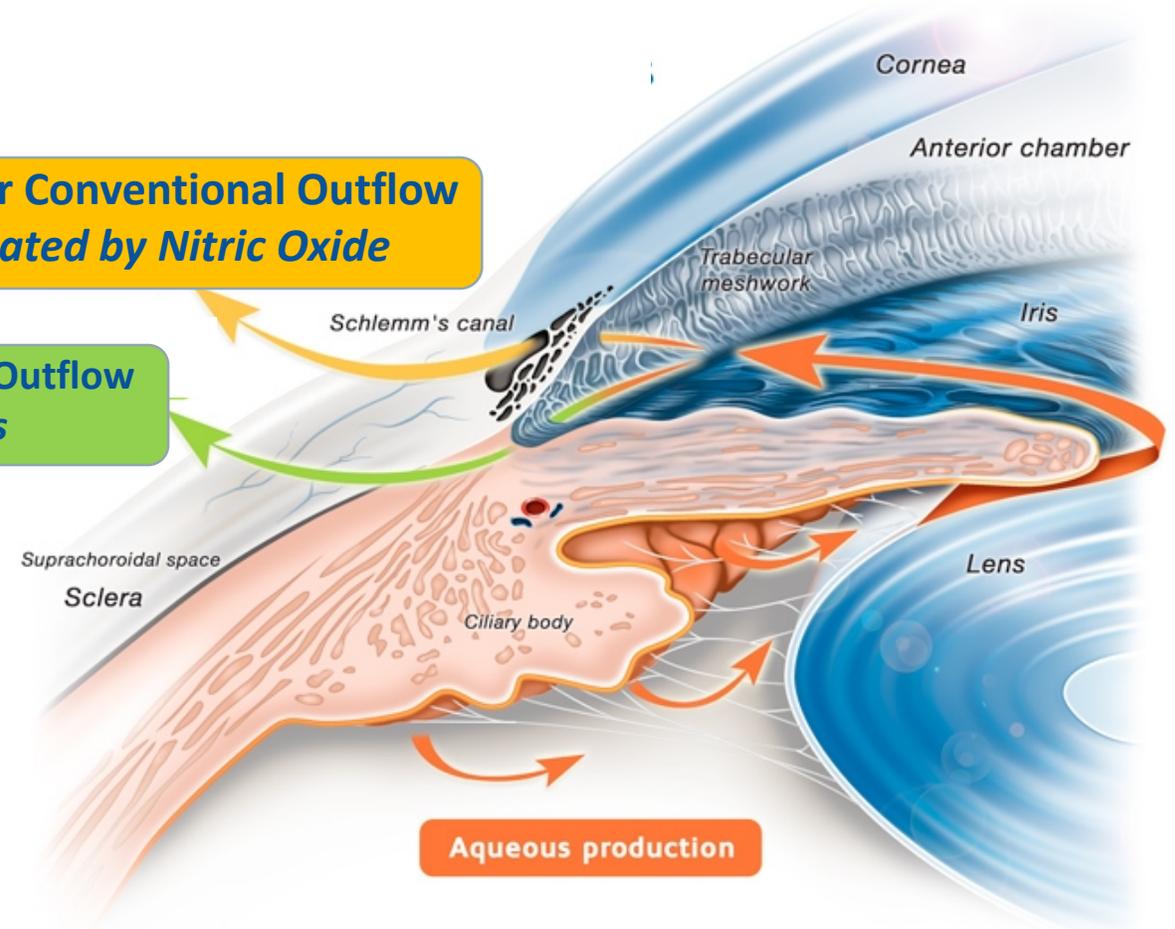
LJ Ignarro

F Murad

**Endogenous cell-
signaling molecule**

**Primary or Conventional Outflow
Stimulated by Nitric Oxide**

**Secondary or Uveoscleral Outflow
Stimulated by PGAs**

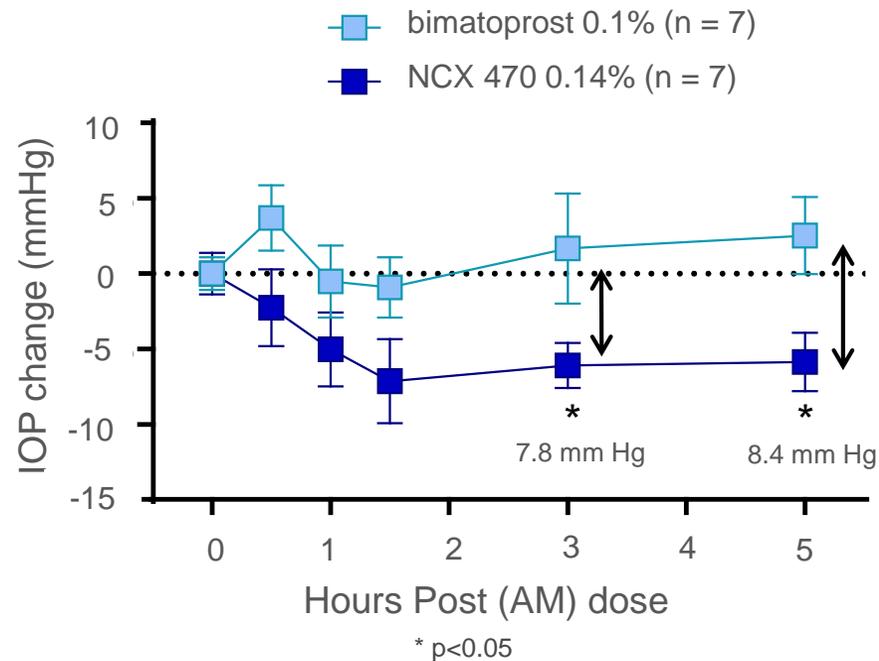


NO in ophthalmology

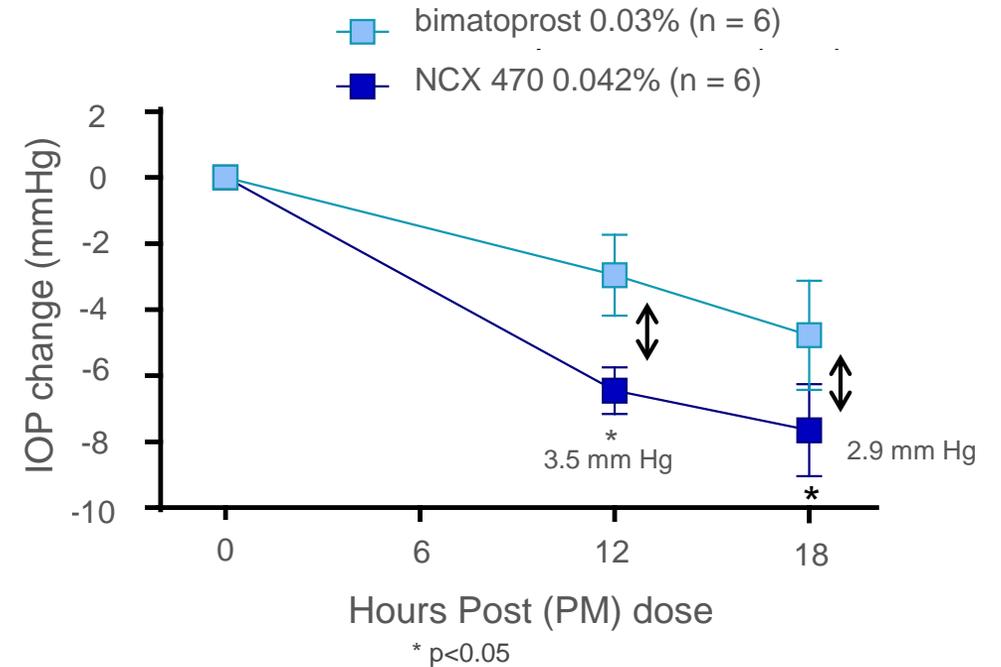
Present in ocular tissues, NO decreases IOP by increasing the outflow of fluid through the primary outflow pathway

NCX 470 Demonstrated Robust IOP Lowering Effect in Vivo

Up to 8.4 mmHg IOP Lowering Due to NO Alone in a Model Poorly Responsive to Bimatoprost
 Transient hypertonic (5%) saline-induced ocular hypertensive rabbits¹



Up to 3.5 mmHg Greater IOP Lowering vs. High Strength Bimatoprost (0.03%)
 Laser-induced ocular hypertensive non-human primates¹



The effect on IOP of nitric oxide in NCX 470 can be clearly demonstrated as separate, complementary and additional to that of the PGA component in animal models

NCX 470 - Phase 2 Clinical Study Initiated August 1st, 2018

Over 85% of Patients Enrolled in the Study (April 18th, 2019)

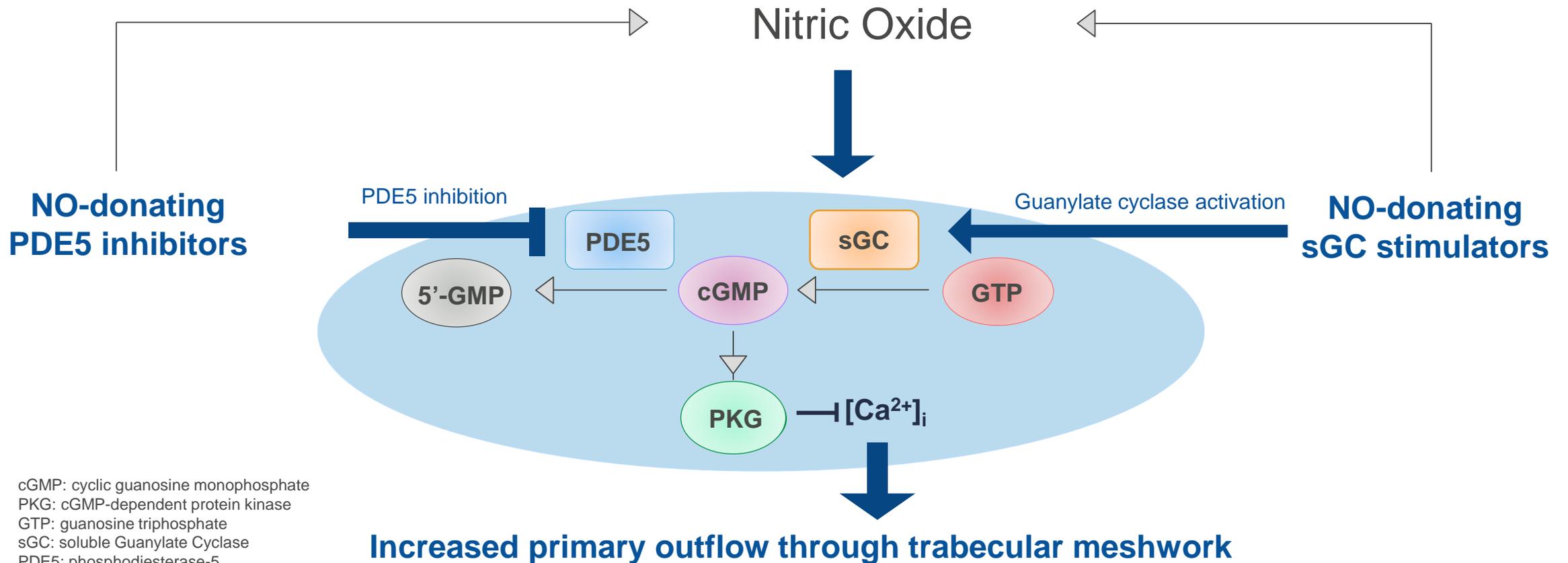
- **Phase 2**, multi-center, double-masked, 28-day, parallel group, dose-response study
- Evaluating the **efficacy and safety** of NCX 470 **compared to latanoprost** 0.005% for IOP lowering in patients with open-angle glaucoma or ocular hypertension
- **420 patients** to be randomized at clinical sites across the U.S.
- Primary efficacy endpoint: **reduction from baseline in mean diurnal IOP** after 28 days of treatment
- Study powered for both **non-inferiority** and **superiority** comparison to latanoprost
- Overall objective: **identification of the appropriate dose** of NCX 470 **to be advanced into Phase 3**
- **Top-line results expected in Q4 2019**

Nicox is committed to using a prostaglandin analog as the active comparator in Phase 3 studies of NCX 470

Future Generation NO-Donors

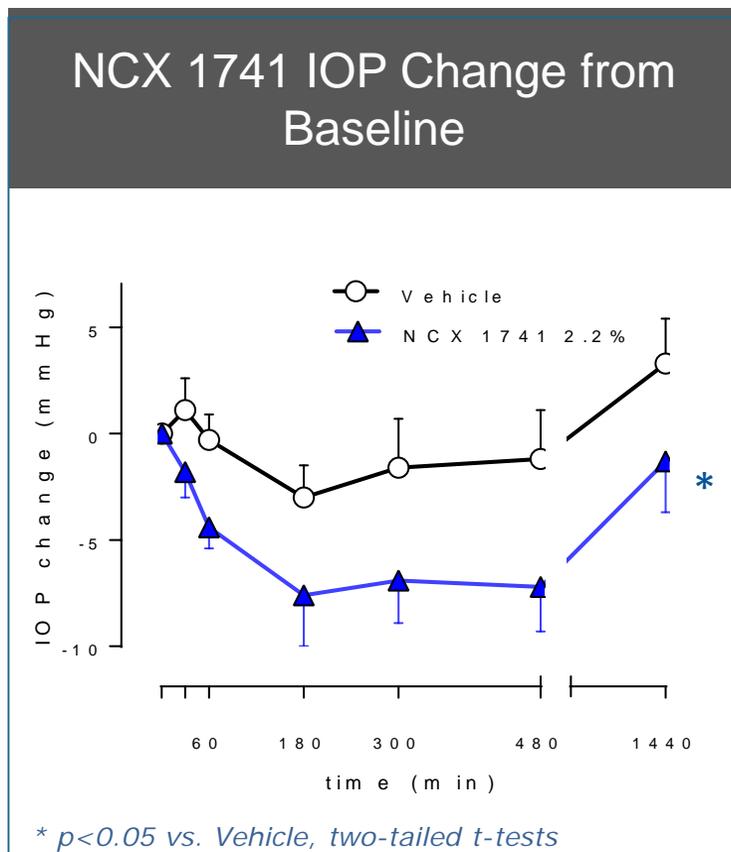
NO-signaling pathway

Enhanced with soluble guanylate cyclase (sGC) stimulators
Prolonged in the presence of phosphodiesterase-5 (PDE5) inhibitors



cGMP: cyclic guanosine monophosphate
PKG: cGMP-dependent protein kinase
GTP: guanosine triphosphate
sGC: soluble Guanylate Cyclase
PDE5: phosphodiesterase-5

NO-Donating PDE5 Inhibitors: a New Class of NCEs for IOP Lowering



- **NO-PDE5 inhibitors:** a new horizon in glaucoma research with a novel **non-PGA chemistry and non-PGA pharmacology**
- Fully **adjunctive to PGA therapy** by targeting exclusively primary outflow
- **Potential for fixed dose combination** products with any PGA
- **Two NCE lead molecules** in formulation optimization towards clinical candidate selection

Robust and sustained IOP lowering effect in non-human primate model of ocular hypertension

NCX 4251 - Phase 2 Clinical Study Initiated March 18th, 2019

Designed to select the dose(s) of NCX 4251 for next stage of development

- **Phase 2**, multi-center, randomized, double-masked, 14-day, placebo-controlled, dose-escalation study
- Evaluating the **safety and tolerability** of NCX 4251 **compared to placebo** in patients with acute exacerbations of blepharitis
- **30 patients** to be randomized at clinical sites across the U.S.
- Primary objective: **selection of the dose(s) for the next larger Phase 2b study**
- **Directly targeting eyelid margin**, where blepharitis disease originates, **via a novel route of delivery**
- **Top-line results expected in Q4 2019**



NCX 4251 represents an opportunity to provide a more efficacious and better tolerated therapy for acute exacerbations of blepharitis compared to the currently available treatments

VYZULTA® - Commercialized in the U.S. by Partner Bausch + Lomb

- First eye drop **approved in 20 years** with a **novel approach** to IOP lowering
- A novel IOP lowering agent **with a dual mechanism of action**
- Proven **IOP lowering up to 7-9 mmHg**
- **Approved in Canada** as of January 2019

Exclusive worldwide license to Bausch + Lomb

Milestones	\$22.5 million in net payments received to date Up to \$150 million net¹ in potential future milestones Majority of remaining milestones based on achievement of target sales
Royalties	6% to 12% net¹ based on global sales of VYZULTA
Exclusivity	U.S. patent extension possible from 2025 to 2030²

ZERVIAE™ - U.S. Commercial Launch by Partner Eyevance Planned in Summer 2019



The first and only topical ocular formulation of cetirizine, indicated for the treatment of ocular itching associated with allergic conjunctivitis

- **Same active ingredient as ZYRTEC®¹** with established systemic efficacy and safety profile in oral formulations resulting from 20 years² of use
- More than **75 million people** suffer from allergic conjunctivitis in the U.S.
- **U.S. topical ocular anti-allergic market approximately \$600 million³**
- **Branded Rx products** represent **~70% market share³**

Milestones	\$6 million upfront received, with up to \$3 million in near-term manufacturing milestones ⁴ and \$37.5 million in potential future sales milestones (of which \$30 million is in milestones triggered by annual sales targets of \$100 million and above)
Royalties	8% to 15% based on future U.S. sales of ZERVIAE
Exclusivity	U.S. patents to 2030 and 2032, Japan patents to 2030

Ongoing discussions for additional licensing agreements ex-U.S.

Strategic Collaborations in China with Ocumension Therapeutics



Validated Partnerships with Ophthalmology Company Funded by a Leading Global Healthcare Investment Fund

NCX 470 for glaucoma

Milestones	<ul style="list-style-type: none"> One-time upfront payment of €3 million Further €2.5 million at the initiation of a Phase 3 study by Nicox with NCX 470 outside the agreed territory. Additional milestones payment up to €14.5 million linked to Ocumension's progress with NCX 470 development, up to and including approval. Up to €16.25 million split over three separate sales milestones associated with potential sales in the territory of up to €200 million
Royalties	<ul style="list-style-type: none"> 6% to 12% on sales

ZERVIATE™ for allergic conjunctivitis

Milestones	<ul style="list-style-type: none"> Development and sales milestone payments of up to €17 million
Royalties	<ul style="list-style-type: none"> 5% to 9% on sales

Financial Highlights

Key Capitalization Overview

Cash & Cash Equivalents¹	€23.5 million
Outstanding shares²	~29.9 million
Fully diluted shares³	~31.3 million
Free float	~97%

Additional Key Statistics

- **Lean organization with 34 employees** in France (Headquarters), Italy (Research Center) and U.S. (Development Center)
- **Debt facility for up to €20 million from Kreos Capital** – 1st tranche of €8 million, 2 other tranches optional at Nicox's sole discretion
- **Minority shareholder in VISUfarma**, a private pan-European ophthalmic specialty pharmaceutical company
- **Future potential royalty from U.S naproxcinod partnership**

A Look into Our Future

Our goal is to become a specialty ophthalmology company driven by its internal R&D pipeline



VYZULTA[®] and ZERVIATE[™] will generate significant and increasing worldwide revenues to support and boost our growth



NCX 470 and NCX 4251 will be the first drugs to be marketed in the U.S. directly by a Nicox commercial organization



Our successful long term growth will be fueled by our innovative R&D pipeline allowing continued organic growth and strong potential for licensing and M&A

Innovative Solutions to Help Maintain Vision and Improve Ocular Health

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