Nicox Corporate Presentation

An international ophthalmology company developing innovative solutions to help maintain vision and improve ocular health

November 22, 2022



Forward-Looking Statements

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Risk factors which are likely to have a material effect on Nicox SA's business are presented in the 3rd chapter of the "*Document d'Enregistrement Universel, rapport financier annuel et rapport de gestion* 2021" filed with the French Autorité des Marchés Financiers (AMF) on April 29, 2022 under number D.22-0392, in its first amendment filed with the AMF under number D.22-0392-A01 on 19 May 2022 and in its second amendment filed with the AMF under number D.22-0392-A02 on November 22, 2022 available on Nicox SA' website (www.nicox.com).

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Driving Innovation in Ophthalmology, Led by NCX 470 & an Experienced Team

Differentiated pipeline with recent, positive NCX 470 pivotal Phase 3 clinical trial results

Lead asset NCX 470, with a potentially differentiated profile targeting glaucoma, leverages Nicox's proprietary Nitric Oxide (NO) donating research platform. Positive topline results from the first Phase 3 trial (Mont Blanc) announced October 31, 2022. Potential retinal benefit seen in nonclinical models¹ Experienced Leadership, Board and Advisors with expertise to drive successful outcomes

Experienced team well positioned to bring NCX 470 to approval and to advance and build the pipeline to deliver future growth Cash position enhanced by global partnerships and outlicensed commercial products

Cash balance of €31.0 million² expected³ to fund operations until mid-May 2024

Current and potential future revenue and value from global partnerships



J Ocul Pharmacol Ther. 2022, 38: 496-504

2. Non-audited figures estimated based on cash at September 30,2022 and including proceeds of the financing announced on November 22, 2022

3. Based on the development of NCX 470 exclusively

Broad Global Leadership Experience





Andreas Segerros Chief Executive Officer

PHARMACIA





Sandrine Gestin VP, Finance

Doug Hubatsch EVP, Chief Scientific Officer



Emmanuelle Pierry General Counsel & Head, Legal

U NOVARTIS

Alcon



Former member of

the Paris Bar



Gavin Spencer EVP, Chief Business Officer & Head, Corporate Development







Board Bringing Extensive Experience in Ophthalmology and Pharmaceuticals



JEAN-FRANÇOIS LABBE Chairman of the Board



Hoechst Marion Roussel



LES KAPLAN Director æiex ≪∷Allergan



MICHELE GARUFI Director





LAUREN SILVERNAIL Director REVANCE[®]



ADRIENNE GRAVES Director Alcon Santen



LUZI VON BIDDER Director





Unique Combination of Competencies

Capable of bringing NCX 470 to approval and driving future growth





Corporate, Finance and Legal team have completed multiple transactions, restructuring and financing



International R&D Management with deep ophthalmology experience



World-recognized Key Opinion Leaders on the Clinical Advisory Board







Novel molecule for intraocular pressure lowering, the leading cause of glaucoma

Positive pivotal Phase 3 topline results from the Mont Blanc trial announced October 31, 2022

Large and established market¹:

\$5.9 billion worldwide

\$1.3 billion prostaglandin analog market in the United States

First non-combination product to demonstrate statistical non-inferiority to a prostaglandin analog in a pivotal trial, thereby meeting the efficacy requirements for approval in the United States



NCX 470 Leads a Differentiated Ophthalmology Pipeline



1. In addition to our Chinese partner, the Company is actively looking for commercial partners in the U.S. and Japan, to maximize the value of NCX 470. The ongoing Denali trial is not financed beyond the current cash runway of mid-May 2024. New Phase 3b clinical trials and nonclinical studies concerning the dual mechanism of action and the potential beneficial effects of NCX 470 on the retina are planned to report results in the next 12 to 18 months which may strengthen the therapeutic profile of NCX470 2. Planned costs of nonclinical activities on NCX 1728 are not significant

3. The net book value of NCX 4251 was decreased to zero (reduction of €15.1 million in 2021 and €11.0 million in H1 2022) in the United States due to the additional costs and timings associated with the change in indication, followed by the decision to outlicense the product

4. The net book value of ZERVIATE (€26 million) corresponds mainly to the value of the asset allocated to the Chinese territory, for which the rights were granted to the partner Ocumension. There was an impairment (€12.7 million) to the value in the United States in 2021 taking into consideration changes in the U.S. market for topical anti-allergics

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NCX 470

Leveraging the potent intraocular pressure-lowering effects of nitric oxide and prostaglandin analogs for potentially differentiated treatment in glaucoma



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Glaucoma Snapshot

Elevated intraocular pressure (IOP) contributes to irreversible optic nerve damage, leading to progressive vision loss



As published in the landmark EMGT study "...each mmHg of decreased IOP was related to an approximately 10% lowering [of risk of vision loss progression]"¹



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Unmet Medical Need for Glaucoma Treatment

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Despite having well established first line therapies, including the standard of care, latanoprost, patients do not react to glaucoma medications in the same way, and therefore eye care professionals need multiple treatment options

40% of patients do not achieve their target IOP on existing monotherapies¹ requiring eye care professionals to adjust or change the medication used Many patients require >1 medication which leads to compliance issues^{2,3} Tolerability issues with some medications lead to discontinuations and/or compliance issues⁴

Kass et al, Delaying treatment of ocular hypertension: the ocular hypertension treatment study. Arch Ophthalmol, 2010; 128:276-287

- 2. Robin AL et al, Does adjunctive glaucoma treatment therapy affect adherence to the initial primary therapy? Ophthalmology. 2005; 112:863-868
- 3. Robin et al, Adherence in glaucoma: Objective measurements of once-daily and adjunctive medication use. Am J Ophthalmol. 2007;144:533-540

4. Beckers HJM et al. Side effects of commonly used glaucoma medications: comparison of tolerability, chance of discontinuation, and patient satisfaction. *Graefe's Archive for Clinical and Experimental Ophthalmology* 2008;246(10):1485-90



NCX 470 Targets the Two Key Outflow Pathways for IOP Lowering

Proven dual mechanism of action¹



Nonclinical optic nerve/retinal damage models demonstrate potentially beneficial retinal effects³

- 2. PGAs = Prostaglandin Analogs
- 3. J Ocul Pharmacol Ther. 2022, 38: 496-504



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Positive Topline NCX 470 Mont Blanc Results Released October 31, 2022 Phase 3 program intended to support planned U.S. & China NDA submissions

Designed to demonstrate safety and efficacy of NCX 470 0.1% vs latanoprost 0.005%, defined by intraocular pressure reduction from time-matched baseline at pre-established time points

MONT BLANC: Primary objective of noninferiority achieved

N=691

56 clinical sites in the U.S. & one site in China Adaptive study design selected the 0.1% dose for the duration of the trial

Second efficacy objective, statistical superiority to latanoprost, was not achieved

NCX 470 was statistically superior to latanoprost in intraocular pressure reduction from baseline at 4 of the 6 timepoints, and numerically greater at all 6

DENALI: Enrolling subjects

N=~670

~60 clinical sites in the U.S. & China
Includes a 12-month safety extension
Jointly conducted & equally financed with Chinese partner Ocumension Therapeutics
Topline results expected in 2025



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Mont Blanc Phase 3 Efficacy Trial Design¹

Designed to evaluate NCX 470 vs. established therapy, latanoprost

Randomized, controlled, double-masked, parallel design trial. Patients with open angle glaucoma or ocular hypertension were randomized 1:1 to once-daily treatment with NCX 470 0.1% or latanoprost 0.005%

Primary Endpoint:

Mean intraocular pressure reduction from time-matched baseline at 8 AM and 4 PM at the week 2, week 6 and month 3 visits

Enrollment:

The trial enrolled 691 patients across all arms



Baseline Characteristics, Demographics and Disposition¹

	NCX 470 0.1% N = 328	Latanoprost 0.005% N = 333
Mean Diurnal Baseline (8am+4pm) IOP, mmHg, Study Eye (SD)	26.9 (2.04)	26.8 (2.02)
Gender , n (%) Female Male	200 (61.0%) 128 (39.0%)	188 (56.5%) 145 (43.5%)
Age, Years (SD)	63.6 (10.12)	62.7 (11.73)
Completed the Study	314 (95.7%)	316 (94.9%)
Discontinued Prior to Study Completion	14 (4.3%)	17 (5.1%)
Reasons for Discontinuation		
Adverse Event	8 (57.1%)	6 (35.3%)
Lost to Follow-up	1 (7.1%)	4 (23.5%)
Physician Decision	0	0
Sponsor or IRB Decision	1 (7.1%)	2 (11.8%)
Protocol Violation	0	1 (5.9%)
Withdrawal by Subject	3 (21.4%)	3 (17.6%)
IOP greater than 36 mmHg	0	0
Other	1 (7.1%)	1 (5.9%)



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NCX 470 0.1% IOP Lowering Compared to Latanoprost 0.005%

Significant, sustained IOP-lowering effects



IOP-lowering effect from baseline was 8.0 to 9.7 mmHg for NCX 470 0.1% vs. 7.1 to 9.4 mmHg for latanoprost (reduction from baseline in time-matched IOP at 8 AM and 4 PM across the week 2, week 6 and month 3 visits)

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NCX 470 0.1% IOP Lowering Compared to Latanoprost 0.005%

NCX 470 0.1% achieved non-inferiority vs. latanoprost 0.005%



Non-inferiority criteria: The upper limit of all 6 95.1% confidence intervals were required to be \leq 1.5 mmHg and at least 4 of 6 were required to be \leq 1.0 mmHg

 Denotes statistically significant differences vs latanoprost (p<0.049)

NCX 470 0.1% demonstrated an IOP lowering effect greater than latanoprost 0.005% of up to 1.0 mmHg



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NCX 470 Topline Results Demonstrate Robust Efficacy and Safety¹ All comparisons are based on NCX 470 0.1% and latanoprost 0.005%

IOP-lowering effect from baseline was 8.0 to 9.7 mmHg for NCX 470 vs. 7.1 to 9.4 mmHg for latanoprost

Measured by a change from baseline in time-matched IOP at 8 AM and 4 PM across the week 2, week 6 and month 3 visits

Statistical non-inferiority was met vs. latanoprost in the primary efficacy analysis

The upper limit of the 95.1% confidence limit on the difference in the treatment effect between NCX 470 and latanoprost in change from baseline in time-matched IOP to the follow-up visits (week 2, week 6, and month 3) was \leq 1.5 mmHg at 6 of 6 timepoints and \leq 1.0 mmHg at 6 of 6 timepoints

NCX 470 failed to meet statistical superiority to latanoprost in a pre-specified secondary efficacy analysis of timematched change from baseline IOP. NCX 470 was **numerically superior** to latanoprost at all time points and statistically significant (p<0.049) at 4 of 6 timepoints

NCX 470 was well tolerated

- the most common adverse event was ocular hyperemia in 11.9% of the NCX 470 patients vs. 3.3% of latanoprost patients
- · there were no ocular serious adverse events and no treatment-related non-ocular serious adverse events
- 4.3% of patients on NCX 470 discontinued compared to 5.1% on latanoprost

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Positive NCX 470 Phase 3 Results a Milestone for Nicox

Novel dual action compound designed to leverage PGA and NO for a differentiated profile

NCX 470 was designed to leverage both PGA and NO mechanisms of action to safely and effectively deliver a differentiated profile in IOP lowering for patients with openangle glaucoma or ocular hypertension NCX 470 reduced IOP by 8.0 to 9.7 mmHg in the 691patient Mont Blanc Phase 3 trial

NCX 470 was statistically noninferior to latanoprost

The secondary efficacy objective, statistical superiority to latanoprost, was not achieved, however, NCX 470 was statistically superior at 4 out of the 6 timepoints

Well tolerated with no ocular serious adverse events and no treatmentrelated non-ocular serious adverse events Mont Blanc Phase 3 results may bring NCX 470 closer to U.S. approval

First non-combination product to demonstrate statistical noninferiority to a prostaglandin analog in a pivotal trial

This trial therefore met the efficacy requirements for approval in the United States

Phase 3 program designed to support NDA submission in U.S. and China



Retinal Benefits: A Potential Differentiator for NCX 470?

Elevated IOP is the main risk factor in glaucoma, however a variety of IOP-independent risk factors, including ischemia, contribute to damage of the optic nerve head and the retina, ultimately causing vision loss

Initial exploratory studies generated encouraging results Exploratory nonclinical studies in a well-defined model of ischemia/reperfusion that results in injury to the optic nerve investigated the potential protective effects of NCX 470 on the retina and the optic nerve head

The results¹ suggest that NCX 470 improves ocular perfusion and retinal function in damaged eyes compared to vehicle and may therefore have protective properties

Next Steps

Nonclinical studies and targeted clinical trials are planned to further explore NCX 470's dual mechanism of action and potential benefits on the retina, beyond its IOP lowering properties



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Why Nitric Oxide Could Generate Retinal Benefits





NCX 470 Shows Retinal Cell Protection in a Nonclinical Model¹

Improved ocular perfusion and retinal function in damaged eyes Potential therapeutic properties beyond IOP lowering



p<0.05 vs respective baseline; * p<0.05 vs vehicle at the same time point, Student's t-Test

Detrimental effect of ET-1 on ophthalmic artery hemodynamics was significantly reversed in eyes receiving NCX 470 0.1% bid (p<0.05 vs. vehicle at week 6) Photoreceptor response decline induced by ET-1 was almost completely reversed in eyes treated with NCX 470 0.1% bid (p<0.05 vs. vehicle at week 6)



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Planned NCX 470 Phase 3b Clinical Trials to Evaluate Potential Retinal Benefit

Nitric oxide has been shown to induce vasodilation. NCX 470's ability to lower episcleral venous pressure as well as enhance outflow through the trabecular meshwork will be evaluated in a clinical trial

Retinal blood vessel density will be studied in a separate clinical trial using Optical Coherence Tomography (OCT)-angiography to fully understand the potential effects on retinal blood flow

Together, these trials are designed to validate NCX 470's dual mechanism of action in humans and potentially demonstrate some of the beneficial effects on the retina that have been observed in nonclinical models.



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Commercial Landscape of the United States Glaucoma Market

Mont Blanc Phase 3 results 8-9.7 mmHg IOP reduction

Non-inferior to latanoprost

NCX 470 Profile¹

- Whilst statistical superiority was not met, NCX 470 was numerically up to 1.0 mmHg better than latanoprost at certain timepoints
- NCX 470 was statistically superior to latanoprost at 4 of 6 timepoints
- · Good tolerability

A space in the market

~\$200 million - estimated potential peak net sales in the U.S. for a product with 1.25 mmHg superior to latanoprost³

~\$100-150 million - estimated potential peak net sales range for recently launched branded products in the PGA market² **Potential growth levers**



Data to be developed supporting non-IOP benefits

Commercial strength of the U.S. marketing partner is required⁴

- 2. Management estimates based on IQVIA Forecast Link. Data on sales of recently launched branded products in the PGA market using a 55% gross-net calculation
- 3. Nicox sponsored market research 2019 and 2021 (the forecast includes estimations about the future growth of the market and assumes an appropriate level of reimbursement is available)

4. For new entrants, including NCX 470 (if approved), obtaining reimbursement and getting on the formularies are critical elements to market access and successful commercialization and [significant] commercial investment by a strong marketing partner is required.

NCX 1728

Novel class of molecules for retinal conditions



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NCX 1728: Lead Compound in a New Class of NO-donating Molecules

Combining NOrelease with PDE5 Inhibition MOA* for this novel class of molecules is based entirely on NO-mediated activity NO-mediated effects are enhanced and prolonged by concomitant phosphodiesterase-5 (PDE5) inhibition within the same molecule

Potential in retinal conditions

NO has a role in ocular perfusion which may be beneficial in a number of orphan retinal conditions for which there is no standard treatment

Nonclinical program focused on evaluating MOA

Nonclinical studies underway to evaluate the mechanism of action in models of orphan retinal conditions



NCX 4251

Novel treatment with unique mode of application in dry eye disease



NCX 4251: Novel Approach to Dry Eye Disease

Novel corticosteroid presentation leverages Nicox's unique formulation expertise Novel, patented ophthalmic nanocrystal suspension of fluticasone propionate, a well-established corticosteroid. Fluticasone has 10x affinity for the glucocorticoid receptor vs. dexamethasone, commonly used in ophthalmology

Planned to be the first topical ophthalmic fluticasone product, a two-week, once-daily treatment leveraging Nicox's proprietary formulation technology

Targeting dry eye disease, a \$3.4 billion prescription market in the U.S. Eye Care Professionals require improved short-term treatment for flares and bridging to chronic therapy

Unique delivery device applies drug directly to the eyelid margin, potentially reducing steroid sideeffects

Phase 2 trial supports potential clinical utility in dry eye disease Post-hoc analysis of 224-subject Phase 2b Mississippi trial showed a statistically and clinically significant reduction in dry eye symptoms versus placebo

Nicox reached alignment with U.S. FDA on a 505(b)(2) development path for NCX 4251 and is currently looking for partnerships outside of China to advance development of this program



Mississippi: Post-Hoc Results Puts Dry Eye Disease in Sight



Unique eyelid margin application designed to minimize corticosteroid-induced ocular adverse events





Reduction from baseline in eye dryness score¹ in patients with inferior corneal fluorescein staining score of ≥ 2

Phase 2b Mississippi² Trial Summary

The trial evaluated NCX 4251 in patients with acute exacerbations of blepharitis. Topline results of the trial did not meet primary endpoint of difference between NCX 4251 and placebo in the proportion of patients with complete cure of eyelid redness, debris, and discomfort

Positive post-hoc results from the Mississippi Phase 2b trial suggest NCX 4251 may be effective in dry eye disease. Patients with a baseline score of \geq 2.0 (on a scale of 0 to 4) for fluorescein staining demonstrated a statistically significant difference in change from baseline vs. placebo for eye dryness score and several other symptoms

NCX 4251 was found to be safe and well tolerated over 14 days with no serious adverse events (all events in the NCX 4251 arm were mild)





. Mississippi: U.S. Multi-Center, Randomized, Double-Masked, Placebo-Controlled, Phase 2b Study Evaluating the Safety and Efficacy of NCX 4251 Ophthalmic Suspension, 0.1% QD for the Treatment of Acute Exacerbations of Blepharitis, ClinicalTrials.gov Identifier: NCT04675242

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Glaucoma: An established \$5.9Bn worldwide, \$2.9Bn U.S. market¹ Approximately 3 million patients in the U.S. with open angle glaucoma²

First line, prostaglandin-based therapies represent a \$1.3 billion opportunity in the U.S. alone¹ 40% of patients on existing monotherapies fail to reach target IOP, risking disease progression and vision loss

Positive Phase 3 results are a major milestone for Nicox First Phase 3 trial demonstrated non-inferiority of NCX 470 to latanoprost³

Statistical superiority to latanoprost was not achieved. However, NCX 470 was statistically superior to latanoprost in IOP reduction from baseline at 4 of the 6 timepoints, and numerically greater at all 6 timepoints

Next Steps on the path to NDA submission

Complete analysis of the Mont Blanc trial data

Complete enrollment in the ~670 subjects/~60 sites (U.S. & China) Denali trial

Denali topline results expected after 2024

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https://www.cdc.gov/features/glaucoma-awareness/index.html

Nicox Press Release 31 October 2022

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U.S. Glaucoma Clinical Advisory Board with Leading Experts

DR. ROBERT D. FECHTNER, MD, CHAIRMAN

Professor and Chair of the Department of Ophthalmology at SUNY Upstate Medical University, Syracuse, NY

DR. SANJAY G. ASRANI, MD

Professor of Ophthalmology at Duke University in Durham, North Carolina, and Director of the Duke Eye Center of Cary and the Duke Glaucoma OCT Reading Center

DR. STEVEN MANSBERGER, MD MPH

Vice-Chair, Senior Scientist, and Director of Glaucoma Services and Ophthalmic Clinical Trials for the Devers Eye Institute in Portland, Oregon. Clinical Professor of Ophthalmology at Oregon Health Science University

DR. TOM WALTERS, MD

President of Texan Eye P.A. and Medical Director of Eye LASIK Austin, Advanced Ophthalmic P.A., Keystone Clinical Research

DR. DONALD BUDENZ, MD MPH

Kittner Family Distinguished Professor and Chairman, Department of Ophthalmology, UNC Chapel Hill School of Medicine

DR. ROBERT N. WEINREB, MD

Distinguished Professor and Chair, Ophthalmology, Director of both the Shiley Eye Institute and the Hamilton Glaucoma Center, holder of the Morris Gleich, MD Chair in Glaucoma, and Distinguished Professor of Bioengineering



Partnering Deals Include Potential Future Payments & Royalties \bigcirc

NCX 470	OcuMension 政 旗 维 视	Potentially differentiated treatment for IOP lowering 6% to 12% royalties on future net sales ¹ in China and Southeast Asia Ocumension pays 50% of the Denali Phase 3 clinical trial costs
VYZULTA BA	USCH+LOMB	First eye drop for glaucoma approved in 20 years with a novel approach to reduce IOP Entitled to \$5 million net milestone at \$100 million net sales 6% to 12% net ² royalties on global sales
ZERVIATE	Configuration With Mit Mit Mit Configuration Con	First and only eye drop formulation of cetirizine for allergic conjunctivitis Phase 3 completed by Ocumension ³ in China: Potential for up to \$17.2 million in sales milestones plus 5% to 9% royalties on net sales Commercialized by Eyevance (a wholly-owned subsidiary of Santen Pharmaceutical Co.) in the U.S.
NCX 4251	OcuMension 账 雌 雅 昶	Novel treatment with unique mode of application in dry eye disease Potential for up to \$11.3 million in future milestones plus 5% to 10% royalties on net sales in China by Ocumension ⁴ Company pursuing out-licensing outside China
Ocumension has rights in Chine Net of royalties payable to Pfize		orea at signed with Pfizer in August 2009

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33 З. 4. Ocumension has rights in Chinese markets

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Ocumension has rights in Chinese and SE Asian markets

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Financial Highlights

Cash balance expected to support current operations through mid-May 2024

Estimated Financial Position and Ownership as of September 30, 2022¹ (updated with financing announced 22 Nov)

Cash, Cash Equivalents	€31.0 million (including proceeds from the financing announced 22 Nov)
Long term debt ²	€20.6 million
Cash runway ³	Mid-May 2024
Outstanding Shares ⁴	43.2 million (plus 6.8 million to be issued at closing of financing announced 22 Nov)
Management and Employees Ownership ⁵	<2%
Key Institutional Investors Analyst Coverage	Armistice Capital 13.7% HBM Healthcare Investments (Cayman) 6.0%,
Bryan Garnier	Eric Yoo
Edison Investment Research	Pooya Hemami
H.C. Wainwright	Yi Chen
Kepler Cheuvreux	Arsene Guekam

1. Unaudited results

2. Includes Kreos Capital bond financing agreement (€18.6 million) and a non-dilutive loan facility credit agreement (€2 million) guaranteed by the French state related to the COVID-19 pandemic

3. Based on the development of NCX 470 exclusively

4. Existing outstanding shares as of October 17, 2022

5. To the best of our knowledge, based on issued share capital



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Value-Creating Milestones

In progress

Building a high-value ophthalmology pipeline

October 2022

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NCX 470 Mont Blanc Phase 3 topline results

Throughout 2023

Communication of Mont Blanc results at key ophthalmology congresses 2025

NCX 470 Denali Phase 3 (enrolling); topline results

H1 2023

Initiation of Phase 3b clinical trials evaluating NCX 470 MOA* and potential beneficial effects on the retina

2024

New nonclinical and clinical data on NCX 470 MOA* and potential beneficial effects on the retina



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*MOA = mechanism of action

Completed

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