



Nucorion Pharmaceuticals, Inc.

Introduction

Nucorion Overview

Innovation Focused on the Chinese Market

- US-based company focused on treatments for **liver-related diseases in China**
- Founded in 2015, with technologies from Ligand Pharmaceuticals (Nasdaq: LGND) and contributions from a faculty member of The Ohio State University
 - There is an established heritage of **successful companies based on Ligand technologies** (Retrophin and Viking Therapeutics)
 - Active involvement from scientific advisors that include **world-renowned leaders** in fields of drug discovery and delivery
- Series A funding of \$5 m (July 2016) has yielded **promising preclinical proof-of-concept data** for Nucorion's lead programs
 - Silver River International Investments Ltd. and Ligand invested in the Series A
 - Efficient operation has allowed substantial progress with minimal cash burn
- First and second parts of Series B funding of \$7.5 m (August 2020) designated for US phase 1 clinical development of hepatitis B program and IND-enabling study of HCC program
 - Ji-Bao pharmaceuticals and Ligand participated
 - Ji-Bao licensed China rights of the hep-B lead compound and responsible for the clinical development in China
- Seeking additional **\$10 million** in Series B funding to obtain clinical proof-of-concept for the second program



Liver Diseases in China

A Growing Epidemic



- Due to economic growth, lifestyle/environmental changes, national policies and an aging population, China is the **fastest growing** pharmaceutical market among emerging countries¹
- The liver-related markets that Nucorion's pipeline is addressing are currently at \$2 b USD, and are **expected to double** to nearly \$4 b by 2024²
- Liver diseases in China are poised to create a **global healthcare burden**.
Key market facts³:
 - **~300 million people** are affected by liver diseases in China
 - Liver cancer is **second most common cause of death** in China
 - China accounts for **over half** of liver cancer deaths worldwide
 - Hepatitis B carrier rate in China **now approaching 10%** of population (compared to <1% in US and EU)⁴

¹ Ni, et al. *Globalization and Health*, 2017, 31:21

² Global Data, *Pharma Intelligence*, 2018

³ Wang, et al. *Hepatology*, 2014, 2099-2108

⁴ Yan, et al. *J. Clin. Translational Hepatology*, 2014, 15-22

Nucorion Overview

Cutting-edge LTP™ Technology

- Proprietary liver-targeting prodrug (LTP) technology to generate *best-in-class* novel medicines by **activating drugs in the liver**
 - Technology is designed to increase liver concentration of actives to maximize efficacy and/or potency at site of interest
 - Also reduces drug exposure outside of liver, minimizing side-effects
- Licensed for Nucorion programs from Ligand Pharmaceuticals
- Recent data has demonstrated **clear preclinical superiority** over ProTide technology used in Sovaldi® and VEMSIDY® (Gilead), and Acelarin (NuCana)
 - Data presented at *EASL's International Liver Conference, 2018*



Nucorion Overview

Scientific Advisory Board

- Nucorion benefits from the active involvement from scientific advisors that include **world-renowned leaders** in fields of drug discovery and delivery



Stephen Benkovic, Ph.D.

- National Medal of Science recipient (2010)
- Penn State University (Professor/Chair of Chemistry)
- National Academy of Science member
- Anacor Pharmaceuticals Co-Founder



Jack Szostak, Ph.D.

- Nobel Laureate (2009 Nobel Prize for Medicine)
- Harvard Medical School (Professor of Genetics)
- Howard Hughes Medical Institute (Investigator)
- Massachusetts General Hospital (Distinguished Investigator)
- National Academy of Science member



John Kozarich, Ph.D.

- Ligand Pharmaceuticals (Chairman); Scripps (adjunct professor)
- ActivX Biosciences (Chairman/President)
- Merck & Co. (Head of Research)
- University of Maryland and Yale University (Professor)



Lin Zhi, Ph.D.

- Ligand Pharmaceuticals (VP, Chemistry)
- Over 25 years discovery & development experience
- Co-author of >60 publications
- Inventor on >100 issued patents
- Inventor of LTP™ Technology platform



Zucai Suo, Ph.D. Co-Founder and Chair of Scientific Advisory Board

- Florida State University
- The Ohio State University (Professor of Chemistry & Biochemistry)
- Harvard Medical School (Postdoctoral Fellow)
- Eli Lilly & Co



Nucorion Overview

Robust, Lower-Risk Pipeline

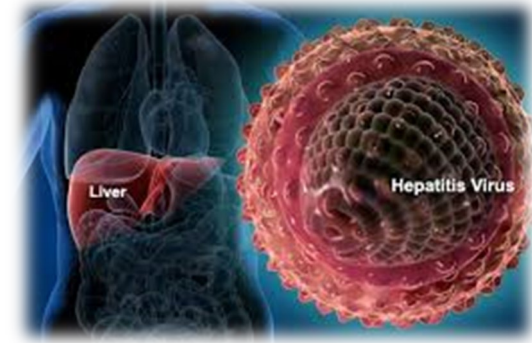
- Multiple **potential *best-in-class*** molecules for an array of liver-related indications
 - All are novel prodrugs of **clinically-validated** actives
 - Prodrug design **avoids off-target safety concerns**

Program Description	Preclinical Development	IND-ready	Phase 1	Upcoming Development Events
NCO-1010 <i>LTP-enabled Nucleotide Analog for Hepatitis B</i>				<ul style="list-style-type: none"> • IND/Phase 1a: Complete • Phase 1b initiation: 2H 2020
NCO-2020 <i>LTP-enabled agent for Hepatocellular Carcinoma</i>				<ul style="list-style-type: none"> • IND: 3Q 2021 • Clinical POC: 2H 2022
NCO-3030 <i>LTP-enabled agents for Solid Tumors</i>				<ul style="list-style-type: none"> • IND: TBD¹ • Clinical POC: TBD²

¹ TBD if the candidate compound distinct from NCO-2020

² Depending upon funding availability

HBV Opportunity



- Established, multi-billion dollar market dominated by Gilead
 - VEMLIDY® (TAF): only new HBV drug in a decade
 - Viread® (TDF) and VEMLIDY® - 2018 Gilead global sales of <\$3b¹
- China situation analysis
 - Hepatitis B carrier rate in China **now approaching 10% of population**²
 - Viread® generic approved in 2017
 - Generic entecavir is the leading HBV drug
 - VEMLIDY® approved 4Q 2018
- Nucorion program (NCO-1010), potentially **best-in-class**
 - Potentially **10-fold more potent** than TAF
 - Potentially **>3-fold more liver-selective** than TAF
- Competitions landscape
 - multiple novel drugs in development, but no immediate success in sight
 - Combo regimen expected with multiple mechanisms, nucleotide-based drug is expected to be the core of treatment

¹ Gilead Q4 2018 Earnings Results Presentation
² Yan, et al. *J. Clin. Translational Hepatology*, 2014, 15-22

HCC Opportunity



- Market approaching a billion USD for small molecule drugs (~\$700m in 2017)¹
 - Market leaders are Nexavar[®] (sorafenib) and Lenvima[®] (lenvatinib), Stivarga[®] (regorafenib), for 2nd line HCC
 - No nucleotide-based drug approved for HCC
- China situation analysis
 - Liver cancer is **second most common cause of death** in China³ and treatment options limited
- Nucorion program – NCO-2020, potentially *first-in-class*
 - Oral delivery of a potent nucleotide agent selectively to the liver for potential use with transartery chemoablation (TACE) before systemic therapies
 - Lowered development and regulatory risks
- Competitive Landscape
 - New kinase inhibitors approved in US and in China⁴ second line with limited efficacy
 - Immuno-oncology (I-O) therapy; Tecentriq approved as 1st line; Opdivo 2nd line
 - I-Os will boost market size, although response rate remains low

¹ Mordor Intelligence, Liver Cancer Market, 2017

² Bayer Annual Report 2017

³ Wang, et al. *Hepatology*, 2014, 2099-2108

⁴ Eisai Major R&D Pipeline, February 2, 2018

Nucorion Overview

Current Status and Plans

- Data obtained post Series A has **clearly validated** the LTP hypothesis
 - Preclinical research goals were achieved on time and under budget
- **Series B investment** will be focused on *Proof-of-Concept* in humans
 - Raising \$17.5 million from US and Asia-based investors (\$7.5 million completed)
 - Positioned to obtain **clinical proof-of-concept** data for two programs addressing billion dollar markets within two years from funding close

Nucorion Overview

An Attractive Investment Thesis

- Potential for a leverageable and **lower-risk drug development platform**
 - Enhancing efficacy and safety profile of validated actives to create *best-in-class* novel medicines for the Chinese market
- **A clear and rapid plan** to clinical *proof-of-concept*
 - Programs quickly approaching or at Clinical Candidate stage
 - Clinical trial designs based on protocols of previously-approved drugs
- **High potential return** for Series B investors
 - Novel, patentable drugs for validated targets, for growing markets with high unmet medical needs
 - Low operating cost structure focuses investment into value-driving events
 - Clinical proof-of-concept for two programs in less than 24 months
 - Potential IPO event in 24-36 months

