

Improving Lives, Offering Hope

Developing New Therapies for Cardiovascular and Neurological Disorders

Corporate Presentation

(ASX:NYR)



January 2020

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Nyrada Introduction and Vision



Intro

 Nyrada listed on the Australian Securities Exchange 16 January 2020 (ASX:NYR)

Vision

To become a high-growth pharmaceutical company, specialising in drug discovery and early-stage development in areas of substantial unmet clinical need, where few (if any) effective or well-tolerated therapies exist

Portfolio

Program 1

A drug to lower cholesterol levels beyond what can be achieved with statins

Program 2

A drug to minimise cell damage associated with brain injury

Program 3

A drug to treat pain associated with nerve injury (sciatica)

Program 1: Cholesterol-lowering drug



Aim

Single-pill treatment for **high cholesterol** (PCSK9i + Statin)

Problem





43 million

ADULTS IN THE US

have high LDL cholesterol and are taking statin medication

US\$317 billion

CARDIOVASCULAR DISEASE COSTS IN THE US

Including healthcare services, medications and lost productivity





Current Standard of Care



Standard Treatment

Statin drugs block the liver's ability to make LDL-cholesterol (Global sales of statin drugs in 2017 estimated to be **US\$19 billion**)

However...

- Many patients are statin intolerant (up to 20%)
- Many patients do not achieve target 'healthy' cholesterol levels (up to 20%)

(Gorcyca K, et al - see slide 7 for more details)

• Approx. half patients have sub-optimal response (<40% lowering) Akyea RK, et al Sub-optimal cholesterol response to initiation of statins and future risk of cardiovascular disease Heart 2019;105:975-981.

Opportunity

Identified need to supplement statin therapy

PCSK9: Beyond Statin Therapy



Cholesterol and PCSK9

PCSK9, a protein found in blood, plays a key role in <u>regulating LDL cholesterol levels</u>

Statins increase PCSK9 blood levels:

This accounts for their failure to work optimally for many patients

PCSK9 inhibitors: If the <u>action of PCSK9</u> is blocked, statins work more effectively, with LDL-cholesterol levels falling an additional 50-60%

Current PCSK9 inhibitors



Two PCSK9-inhibitors came to market in 2015 Effective when used in combination with statins, however ...

- Must be injected every 2-4 weeks, for life
- High cost (approx. US\$5k per year)

Our Cholesterol-Lowering Solution



What's Unique

As far as we are aware, Nyrada is only 1 of 2 companies developing a small molecule PCSK9-inhibitor. Our aim is that our drug will pave the way for a **single pill solution**, for effective lowering of high LDL cholesterol.

- ✓ Benefit of allowing a **lower statin dose in statin-sensitive patients** (PCSK9 inhibitor + statin)
- ✓ Monotherapy treatment in patients unable to tolerate statins (Nyrada PCSK9 inhibitor alone)

Target Drug Profile

- Once-a-day oral tablet incorporating a generic statin
- Safety profile consistent with chronic administration
- Patentable
- Cost effective

Cholesterol-Lowering Drug: Market Overview (US)



Patient and Market Need

Large potential market – Approx. **18 million people** in the U.S. with <u>atherosclerosis</u> who live with <u>elevated LDL levels</u> despite taking <u>maximally tolerated statin therapy</u> — including individuals considered statin intolerant — leaving them at <u>high risk for cardiovascular events</u>.

Source: Gorcyca, K., et al., Prevalence of Atherosclerotic Cardiovascular Disease (ASCVD) and Diabetes Populations in the United States. Journal of Clinical Lipidology, 2015. 9(3): p. 424.

Overview of competitor PCSK9 therapies

Drug Name	Status	Company	Target	Molecule	Delivery
Evolocumab (Repatha)	Marketed	Amgen	PCSK9 inhibitor	Monoclonal	Injectable
Alirocumab (Praluent)	Marketed	Sanofi/Regeneron	PCSK9 inhibitor	Monoclonal	Injectable
Bempedoic acid ± ezetimibe	Phase III	Esperion	ATP citrate lyase inhibitor	Small molecule + combination	Oral
Inclisiran	Phase III	The Medicines Company	PCSK9 siRNA	siRNA	Injectable
Evinacumab	Phase III	Regeneron	ANGPTL3 inhibitor	Monoclonal	Injectable
LY3015014	Phase II	Lilly	PCSK9 inhibitor	Monoclonal	Injectable
AFFITOPE (AT04A)	Phase I	AFFIRIS AG	PCSK9	Vaccine	Injectable
P-21	Preclinical	Shifa Biomedical	PCSK9 inhibitor	Small molecule	Oral
NYX-330	Preclinical	Nyrada Inc.	PCSK9 inhibitor	Small molecule	Oral

Program 2: Brain Injury Drug



Aim

Treatment for Stroke and Traumatic Brain Injury



Stroke

0.8 million

PEOPLE EACH YEAR suffer a stroke in the US



STROKE COSTS

Direct medical costs and indirect costs

US\$34 billion

yearly in the US

Traumatic Brain Injury

2.8 million

PEOPLE EACH YEAR sustain a TBI in the US

TBI COSTS

Direct medical costs and indirect costs

US\$60 billion

yearly in the US

Brain Injury Solution



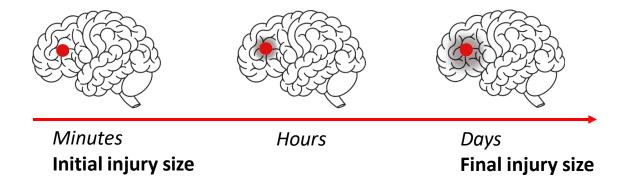
Current Treatment Options

Largely unmet clinical need

- Clot-buster drugs for treatment of ischemic stroke
 (Only suitable for approx. 10% of acute stroke patients due to short (<4.5 hr) therapeutic window)
- Drug treatments for brain injury limited to diuretics, anti-seizure, and coma-inducing drugs

Nyrada Drug

- In the days following brain injury, the area of damage expands, worsening patient outcomes
- Drug aims to prevent damage, limiting injury size and improving patient outcomes



Brain Injury Drug: Market Overview (US)



Stroke	
Stroke emergency room visits per year in the US	Approx. 650k
Patients who present within nominal 12-hour therapeutic window (>75%)	Approx. 490k

Clot-buster Alteplase sales approx. US\$1.2 billion in 2017; cost per patient approx. US\$11k

Traumatic Brain Injury (Moderate to Severe)				
Hospital admissions for TBI per year in the US	Approx. 280k			
Patients who present within nominal 12-hour therapeutic window (>75%)	Approx. 210k			

Total annual US market for brain injury combined: approx. 700k patients*

https://www.ncbi.nlm.nih.gov/pmc/articles/PMC5614785/

Morris DL et al. Prehospital and emergency delays after acute stroke: the Genentech Stroke Presentation Survey: Stroke 2000 Nov;31(11):2585-90 Taylor CA et al. Traumatic Brain Injury—Related Emergency Department Visits, Hospitalizations, and Deaths — United States, 2007 and 2013

Nyrada Inc — 11 — January 20

^{*} However, there is no guarantee that such a market will eventuate and that, even if such a market does eventuate, Nyrada will be able to enter into, and take advantage of, any such a market.

Company Program Status



- Proof-of-concept established for both the cholesterol-lowering and neuroprotection programs
- Lead optimisation well-advanced
 - Composition of matter patents have been lodged (but not yet granted) for both programs
 - Key publication submitted (PCSK9 inhibitor)
- First-in-human Phase I studies target time frames (subject to all necessary regulatory approvals and preclinical results)
 - Cholesterol-lowering program: ready to enter clinic late-2021
 - Neuroprotection program: mid-2022

Board of Directors



Nyrada operates under the direction of a board of international calibre, with a track record in founding and realizing the value of biotech companies



Mr John Moore Non-Executive Chairman John Moore currently serves as Non-Executive Chairman of Trialogics, a clinical trial informatics business. John is a director of Scientific Industries (SCND-OTCQX), a producer of laboratory instruments for the life sciences industry. He is a graduate of Rutgers University.



Dr Graham Kelly Non-Executive Director Dr Graham Kelly is a scientist with 50 years' experience in drug development in both academic and biotechnology sectors. Nyrada,Inc. is the fourth public company founded by Dr Kelly (3 still operational, 2 US-listed). He is the Founder and Executive Chairman of Noxopharm Limited (ASX:NOX), the major shareholder in Nyrada. He holds a PhD in Medical Science as well as degrees in Science and Veterinary Science from The University of Sydney.



Christopher Cox Non-Executive Director Christopher Cox has been a partner at Cadwalader, Wickersham & Taft LLP since 2012 advising clients on a wide array of corporate and financial matters. From February 2016 to March 2019, Chris was seconded to The Medicines Company, where he served as Executive Vice President and Chief Corporate Development Officer and was responsible for business development and strategy. Chris received both his undergraduate degree and J.D. from the University of Missouri, where he was also a member of the Missouri Law Review.

Board of Directors (continued)



Nyrada operates under the direction of a board of international calibre, with a track record in founding and realizing the value of biotech companies



Mr Marcus Frampton Non-Executive Director Marcus Frampton is CIO of Alaska Permanent Fund Corporation (APFC), a \$65 billion sovereign wealth fund for Alaska. He is also a shareholder/Director of Scientific Industries, Inc, a leading manufacturer of laboratory equipment and owner of IP relating to bioprocessing systems. Marcus graduated from UCLA with a degree in Business-Economics and a Minor in Accounting.



Dr Rüdiger Weseloh Non-Executive Director Dr Rüdiger Weseloh joined Merck KGaA, Darmstadt, Germany, as Senior Licensing Manager in 2006 holding positions in BD and is now a Senior Director. He has a university diploma in biochemistry from the University of Hannover and a PhD in molecular neurobiology, obtained at the Center for Molecular Neurobiologyin Hamburg.



Mr Peter Marks
Non-Executive
Director

Peter Marks is currently a Director of Alterity Therapeutics Limited (ASX:ATH and NASDAQ:ATHE), Non-Executive Director of Noxopharm Limited (ASX: NOX) and Non-Executive Director of Fluence Corporation Ltd (ASX: FLC). Peter holds an MBA from the University of Edinburgh, Scotland, a Bachelor of Economics, Bachelor of Laws and a Graduate Diploma in Commercial Law from Monash University, Australia.

Scientific Advisory Board



Nyrada benefits from an international team of experts with deep experience in drug development to advise its board and management

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Prof Gary Housley MSc, PhD Scientia Professor Gary Housley is the Chair of the Nyrada Inc Scientific Advisory Board. Prof. Housley holds the Chair of Physiology and is director of the Translational Neuroscience Facility, School of Medical Sciences at the University of New South Wales, Sydney, Australia



Prof Junichi Nabekura PhD Junichi Nabekura is Professor of Physiology and Neuroscience, and Director of the National Institute of Physiological Sciences (NIPS) in Okazaki, Japan



Prof David Burke MD, DSc, AC Dr David Burke is Professor of Neurology at Royal Prince Alfred Hospital, University of Sydney



Prof Gilles Lambert PhD Dr Gilles Lambert is Professor of Cell Biology at The University of La Réunion Medical School (France) and group leader, Inserm Laboratory of Diabetes & Atherothrombosis of the University Hospital of La Réunion. Since 2004, Dr Lambert has conducted seminal research projects on PCSK9, a major inhibitor of the LDL receptor.



Jim Palmer PhD Dr Jim Palmer brings over 30 years of experience in drug discovery programs targeting oncology, cardiovascular, inflammation, joint and bone disease, and infectious diseases.



www.nyrada.com

- Large and under-served therapeutic areas
- Proof of concept data from cell and animal models
- Right team assembled to execute on strategic objectives

USA

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