21 May 2025

Sydney, Australia

Phase I Clinical Trial Dosing Update #3

Highlights:

- Safety Review Committee (SRC) for Nyrada's Phase I clinical trial has considered cumulative safety and pharmacokinetic data, including from the third dosed cohort.
- SRC raised no issues, enabling the clinical trial to proceed to the next cohort.

Nyrada Inc (ASX:NYR), a drug discovery and development company focused on innovative Transient Receptor Potential Canonical (TRPC) ion channel inhibitors, today provides an update on its Phase I clinical trial.

Clinical Trial

Nyrada has confirmed that the SRC has reviewed cumulative safety and pharmacokinetic data from the first, second, and third cohorts in its clinical trial. No issues were raised, enabling the trial to progress to the fourth cohort for which recruitment has been complete.

Final Phase I trial readouts are expected in the third quarter of the 2025 calendar year. Regular updates will be provided throughout the trial.

Lead Drug Candidate NYR-BI03

Nyrada is developing NYR-BI03, a small-molecule, first-in-class drug for neuroprotection and cardioprotection indications.

In February 2024, Nyrada announced <u>preclinical stroke study results</u> showing that NYR-BI03 achieved a statistically significant neuroprotective effect, rescuing 42% of brain tissue in the penumbra region of treated animals.

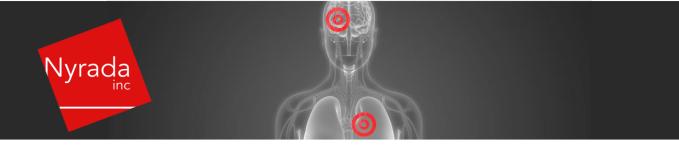
In October 2024, Nyrada announced the results of a <u>preclinical coronary heart disease</u> study, which showed that NYR-BIO3 provided an 86% cardioprotective effect following myocardial ischemic-reperfusion injury, a leading cause of tissue damage when blood flow is restored to the heart after injury. Further <u>supporting efficacy data</u> was provided through echocardiography assessment showing significant improvements in heart function and structure following NYR-BIO3 treatment

In April 2025, Nyrada announced the results of a <u>preclinical traumatic brain injury</u> study which showed that NYR-BI03 provided a statistically significant (p = 0.043) neuroprotective effect following a penetrating traumatic brain injury. This study was undertaken in collaboration with the <u>Walter Reed Army Institute of Research</u> and <u>UNSW Sydney</u>.



In May 2025, Nyrada announced the results of a follow up <u>preclinical coronary heart disease</u> study. This study showed that NYR-BI03 provided 42% cardioprotection when administered continuously for only 3 hours. In addition to protecting the irreplaceable heart tissue, and reducing injury biomarker levels, the incidence of arrhythmia parameters including ventricular fibrillation and ventricular tachycardia, the leading causes of sudden cardiac death, were significantly reduced.

-ENDS-



Appendix 1 - Key Details of NYR-BI03 Phase I Clinical Trial

Protocol Title	A Phase I, Double-Blind, Placebo-controlled, Randomised, First in Human, Dose Escalation Study to Assess the Safety, Tolerability, and Pharmacokinetics of NYR-BI03 in Healthy Participants, When Administered as a 3-hour Infusion	
Primary Endpoints	To evaluate the safety and tolerability of NYR-BI03 in healthy volunteers, when administered as a 3-hour intravenous (IV) infusion	
Secondary Endpoints	To determine the blood pharmacokinetics (PK) of an intravenous dose of NYR-BI03 in healthy volunteers when administered as a 3-hour infusion.	
Blinding Status	Double-blind, placebo-controlled, randomised	
Treatment Method	3-hour intravenous infusion	
Number of Trial Subjects	Up to approximately 40 participants will be enrolled (8 participants per cohort for 5 cohorts)	
Inclusion Criteria	 Informed consent 18 to 50 years of age Male or female Weight 50 to 105 kilograms Healthy as determined by a medical history 	
Exclusion Criteria	 Pregnancy Allergy or hypersensitivity to formulation or ingredients Any evidence of organ dysfunction Liver function or blood clotting tests outside the approved range Drug and alcohol abuse Prescription medications taken within 14 days prior to dosing Psychiatric disorder Blood donation within 12 weeks prior to dosing Vaccination or immunisation within 30 days prior to dosing 	
Trial Location	Scientia Clinical Research The Bright Building Level 5, Corner of Avoca and High Street Randwick NSW 2031 Australia	
Principal Investigator	Dr Christopher Argent Scientia Clinical Research	
Contract Research Organisation	Southern Star Research Level 1, 1 Merriwa Street	
	Gordon NSW 2072 Australia	



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Nyrada Inc. is a biotechnology company focused on the discovery and development of innovative small-molecule therapies, specifically targeting Transient Receptor Potential Canonical (TRPC) ion channels. The company's lead candidate, NYR-BIO3, has shown efficacy in both neuroprotection and cardioprotection, positioning it for a first-in-human Phase I clinical trial. Nyrada Inc. (ARBN 625 401 818) is incorporated in Delaware, USA, with limited liability for its stockholders.

www.nyrada.com

Authorised by Mr. John Moore, Non-Executive Chair on behalf of the Board.

Investors and Media:	Company Secretary:
Dimitri Burshtein	David Franks
T: 02 9498 3390	T: 02 8072 1400
E: <u>info@nyrada.com</u>	E: <u>David.Franks@automicgroup.com.au</u>

Forward-Looking Statements

This announcement may contain forward-looking statements. You can identify these statements by the fact they use words such as "aim", "anticipate", "assume", "believe", "continue", "could", "estimate", "expect", "intend", "may", "plan", "predict", "project", "plan", "should", "target", "will" or "would" or the negative of such terms or other similar expressions. Forward-looking statements are based on estimates, projections, and assumptions made by Nyrada about circumstances and events that have not yet taken place. Although Nyrada believes the forward-looking statements to be reasonable, they are not certain. Forward-looking statements involve known and unknown risks, uncertainties and other factors that are in some cases beyond the Company's control that could cause the actual results, performance, or achievements to differ materially from those expressed or implied by the forward-looking statement.