



Noxopharm Limited ([ASX:NOX](#)) | ASX Announcement | 9 September 2021

Noxopharm Updated Corporate Presentation

Sydney 9 September 2021: Australian clinical-stage drug development company Noxopharm Limited (ASX:NOX) is pleased to provide an updated non-confidential corporate presentation.

The presentation will be used for the upcoming HC Wainwright & Co 23rd Annual Global Investment Conference which is being held as a virtual event from 13-15 September, 2021.

The conference is an annual event for institutional investors, private equity firms, venture capitalists, industry executives and business development executives.

The presentation focuses on the Company's position in the oncology space as having a unique multi-functional drug platform with the potential to assist the annual multi-billion dollar chemotherapy, radiotherapy and checkpoint inhibitor markets and the nascent targeted radioligand therapy market.

A recording of the corporate presentation will be made available on the Noxopharm website late next week. www.noxopharm.com

-ENDS-

Graham Kelly, CEO and Managing Director of Noxopharm, has approved the release of this document to the market on behalf of the Board of Directors.

About Noxopharm

Noxopharm Limited (ASX:NOX) is an Australian clinical-stage drug development company focused on the treatment of cancer and cytokine release syndrome (septic shock).

Veyonda[®] is the Company's first pipe-line drug candidate currently in Phase 2 clinical trialling. Veyonda[®] has two main drug actions – a moderating effect on the ceramide/sphingosine-1-phosphate balance and inhibition of STING signalling. Activity against the former target contributes to its dual-acting oncotoxic and immunomodulatory functions designed to enhance the effectiveness and safety of standard oncology treatments, i.e., chemotherapies, radiation therapies and immune checkpoint inhibitors. Activity against the latter target provides an anti-inflammatory effect, as well as contributing to an anti-cancer action, but also potentially blocking septic shock.



Noxopharm is running comprehensive drug discovery programs in both oncology and inflammation, and is the major shareholder of US biotechnology company, Nyrada Inc (ASX:NYR), active in the areas of drug development for cardiovascular and neurological diseases.

To learn more, please visit: noxopharm.com

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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 Noxopharm about circumstances and events that have not yet taken place. Although Noxopharm 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 (including but not limited to the COVID-19 pandemic) that could cause the actual results, performance or achievements to differ materially from those expressed or implied by the forward-looking statement.



H.C. Wainwright 23rd Annual Global Investment Conference

13-15 September 2021





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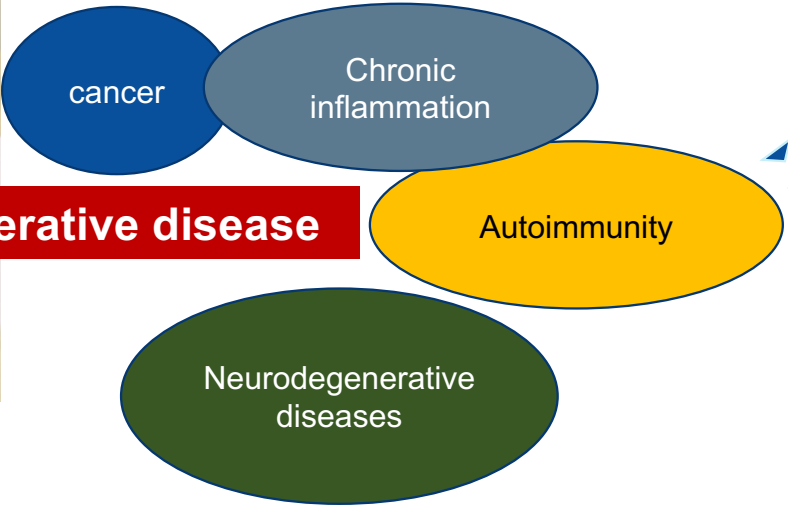
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Veyonda® currently is not approved for use in Australia or any other country.



**Involves multiple mutations,
multiple errors**

Single drugs targeting multiple errors



**Therapies targeting single errors
are restricted in their benefit**

OR

The choice is polypharma

Introducing idronoxil (Veyonda[®])

A novel multi kinase and membrane NAD(P)H-oxidase inhibitor **targeting aberrant protein behaviour**

Primary biochemical targets

Protein thiol-disulfide exchange

Protein phosphorylation, dephosphorylation, dimerisation

Sphingomyelin pathway

Primary biological outcomes

Protein folding

Protein function

Pro-survival/
pro-death signaling

Primary functional outcomes

Anti-cancer effects

Anti-inflammatory effects

Immuno-stimulatory effects

Four R&D Businesses in One Company



Veyonda®

Cancer treatment
enhancement



**Cancer Research
Pipeline**

Cancer growth
factor inhibitors



Veyonda®

Septic shock



pharmorage

Chronic inflammatory
diseases/autoimmune
diseases

Veyonda®

Cancer treatment
enhancement



A revolutionary concept in cancer therapy involving a drug (Veyonda®) with **multiple mechanisms of action** that aims to boost the effectiveness of other standard anti-cancer therapies

Programs



**IONIC - checkpoint
inhibitor therapy**



DARRT - radiotherapy



CEP - chemotherapy



**LuPIN - radioligand
therapy**

Veyonda®

Cancer treatment
enhancement

IONIC

Checkpoint inhibitor therapy

Challenge

- PD-1 inhibitor therapy relies on the presence of effector T-cells to take advantage of drug action
- ~95% of human tumours appear to lack effective immune function ('COLD')

Idronoxil

- Inhibits sphingosine kinase 1, blocking S1P production
- → upregulation of CD4+ and CD8+ T-cells
- → restoration of immune function in tumours ('COLD' to 'HOT')

Rationale

- Sphingosine-1-phosphate (S1P) key regulator of immune cell trafficking
- Most tumours over-express S1P
- High to low S1P gradient between tumour and blood causes egress of T-cells

Phase Ib trial

Veyonda + nivolumab

Cohort 1: 15 patients refractory to PD-1 inhibitor therapy

Cohort 2: 15 patients PD-1 inhibitor therapy naive

Veyonda®

Cancer treatment
enhancement

DARRT

External beam radiotherapy

Challenge

- Low-dose EBRT triggers an immune response within single irradiated tumours
- On very rare occasions, that immune response shifts from a local to a systemic level → **abscopal response**

Idronoxil

- Inhibits autophagy
- Activates immune function

Rationale

- Immune (interferon) response to low-dose EBRT stems from damage to mitochondrial DNA
- That damage is repaired by autophagy
- Blocking autophagy extends the interferon response → **augmented abscopal response**

Phase II DARRT trial

Multi-national (U.S., Australia, France, Hungary)

~100 patients; prostate, breast, lung cancers refractory to standard therapies

Primary end-points: incidence of abscopal responses (RECIST), PFS



Veyonda®

Cancer treatment
enhancement

LuPIN

Radioligand therapy

Challenge

- Effectiveness of targeted radiotherapy by radioligands dependent on expression of antigen target on cancer cells
- ¹⁷⁷Lu-PSMA therapy limited by rate of expression of PSMA on prostate cancer cells

Idronoxil

- Blocks cell division (G₂M) → **greater DNA damage**
- Blocks DNA repair (topoisomerases 1 and 2; PARP1) → **greater cytotoxicity**

LuPIN phase I/II study

- 56 mCRPC patients
- Post-enza/abir and 2x taxanes
- **mOS = 19.7 months**

Rationale

- One strategy is to increase PSMA expression
- Another is to enhance the damage inflicted by the available radioactivity

Veyonda®

Cancer treatment
enhancement

CEP

Chemotherapy

Challenge

High rates of inherent and acquired resistance in most tumour types to cytotoxic chemotherapies due to over-expression of pro-survival mechanisms

Idronoxil

- Blocks external membrane NADH oxidase function → **interferes with protein folding**
- Blocks S1P expression → **reduction in major pro-survival signaling**
- Enhanced ceramide expression → **increased pro-death signaling**
- Increases immune cell recruitment

Rationale

- Priming cytotoxic effects of chemo drugs by reducing pro-survival signaling

Phase Ib CEP-2 trial

IND granted July 2021

Multiple U.S. sites planned

First-line therapy doxorubicin and Veyonda in patients with soft tissue sarcomas

'Helper' growth factor inhibitors



Many cancers, particularly the highly aggressive cancers, co-opt supporting healthy cells to supply growth factors that drive cancer cell growth.

Noxopharm has identified a novel family of drugs with potential to block these signals

Programs



Brain cancer (GBM)



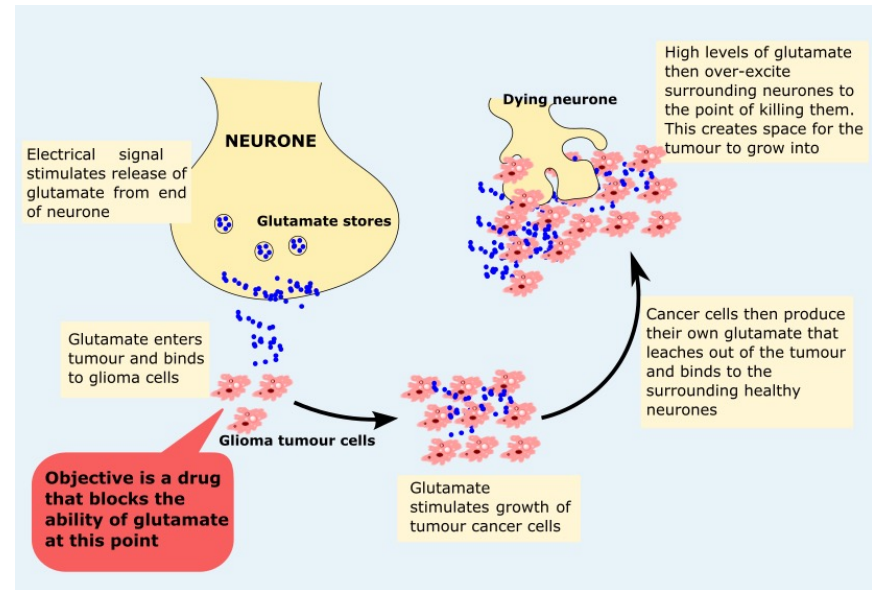
Pancreatic cancer

Brain cancer program

Noxopharm and U.S. National Cancer Institute to Collaborate on Promising New Approach to Treatment of Brain Cancer*

Major discovery by NOX scientists of new family of molecules

- ➔ Killing brain cancer cells directly
- ➔ Blocking 'helper' growth signals



[*ASX Announcement 9 August 2021](#)

Veyonda®

Septic shock

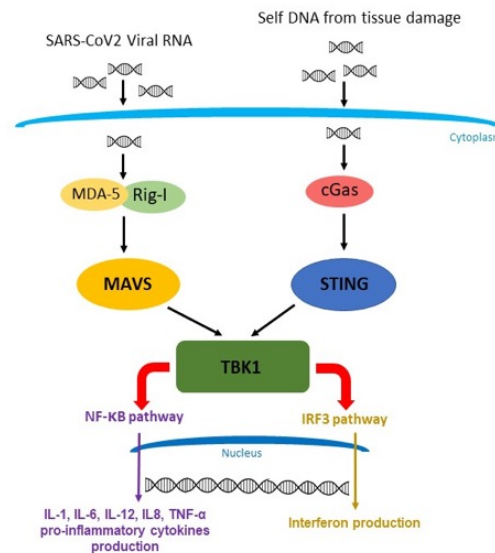


Challenge

To block the inappropriate hyper-inflammatory response to damage caused by the SARS-CoV-2 virus, without blocking a protective anti-viral immune response

Idronoxil

- Potent inhibitor of cGAS-STING /TBK1 signaling pathway → **blocking release of pro-inflammatory cytokines**
- Upregulates NK and T-cell function



Phase I pilot trial completed

- Hospitalised patients with moderate to severe ARDS requiring supplementary O₂
- Veyonda added to SOC
- Treatment well-tolerated
- 37/38 patients recovered/1 death
- Pro-inflammatory cytokines all contained

Phase 2 randomised controlled study proposed for Veyonda to be tested in hospitalized COVID-19 patients with mild hypoxia.

Anti-inflammatory drug offering broad-spectrum cytokine inhibition, but without immune-suppression



pharmorage

Chronic inflammatory diseases/autoimmune diseases

pharmorage

Pharmorage, a collaboration between NOX and Hudson Institute of Medical Research.

Based on identification of a new family of drug compounds with highly selective activity against cGAS-STING/TBK1 signaling*

* ASX: 23 August 2021

Programs



Chronic inflammation



Autoimmunity

STING/TBK1 antagonists now centre of major industry interest as new drug class



Key Metrics

(at 8 September 2021)

Market cap

A\$153m

Share price

A\$0.53c

Issued shares

~288.3m

Cash (at 30/6/21)

A\$26.8 m

Anticipated News Flow (next 6 months)

- ▶ Progress in IONIC-1, DARRT-2 & CEP-2
- ▶ Phase 2 COVID-19 clinical trial update
- ▶ Oncology drug pipeline progress
- ▶ Pharmorage drug discovery progress



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