

# N4 Pharma

## Investor Presentation

MELLO 2018



# Introduction

- Established in 2014, RTO onto AIM May 2017
- Specialist pharmaceutical company with two divisions:
  - Reformulation of existing generic drugs; and
  - Silica nanoparticle delivery system (Nuvec®)
- Lower risk, less costly, quicker to market than traditional drug development
- Lead Generic product, sildenafil MR (more commonly known as Viagra), undertaking proof of concept clinical trials
- Nuvec® research program leading to early collaborations
- Built around a strong IP portfolio in both divisions
- Highly experienced management team

# Business Model

- Develop new versions of existing drugs either already on market (generics) or in development (vaccines)
- License to big pharmaceutical companies for milestones and royalties



- 1**
  - *Unmet patient need*
  - *Global market potential*
  - *Identify & select therapeutic products with side-effects, stability or efficacy issues*
- 2**
  - *Patent reformulations of improved generic*
  - *Improve DNA/RNA delivery using proprietary technology platform*
- 3**
  - *License re-formulated products to global pharma market*
  - *License delivery technology to pharma & biotech for own programs*

# One year on: SP up over 200%



Source: London Stock Exchange

- Commencement of in-vivo research programme for Nuvec®
- First collaboration announced
  - MedImmune to evaluate Nuvec® technology
- Appointment of Andrew Leishman as Head of Nuvec® development
- Commencement of sildenafil human clinical trial
- £784k raised via warrant exercises since 1 January 2018
- Strong cash position through to 2019



# Sildenafil MR

# The Perfect Erectile Dysfunction Product

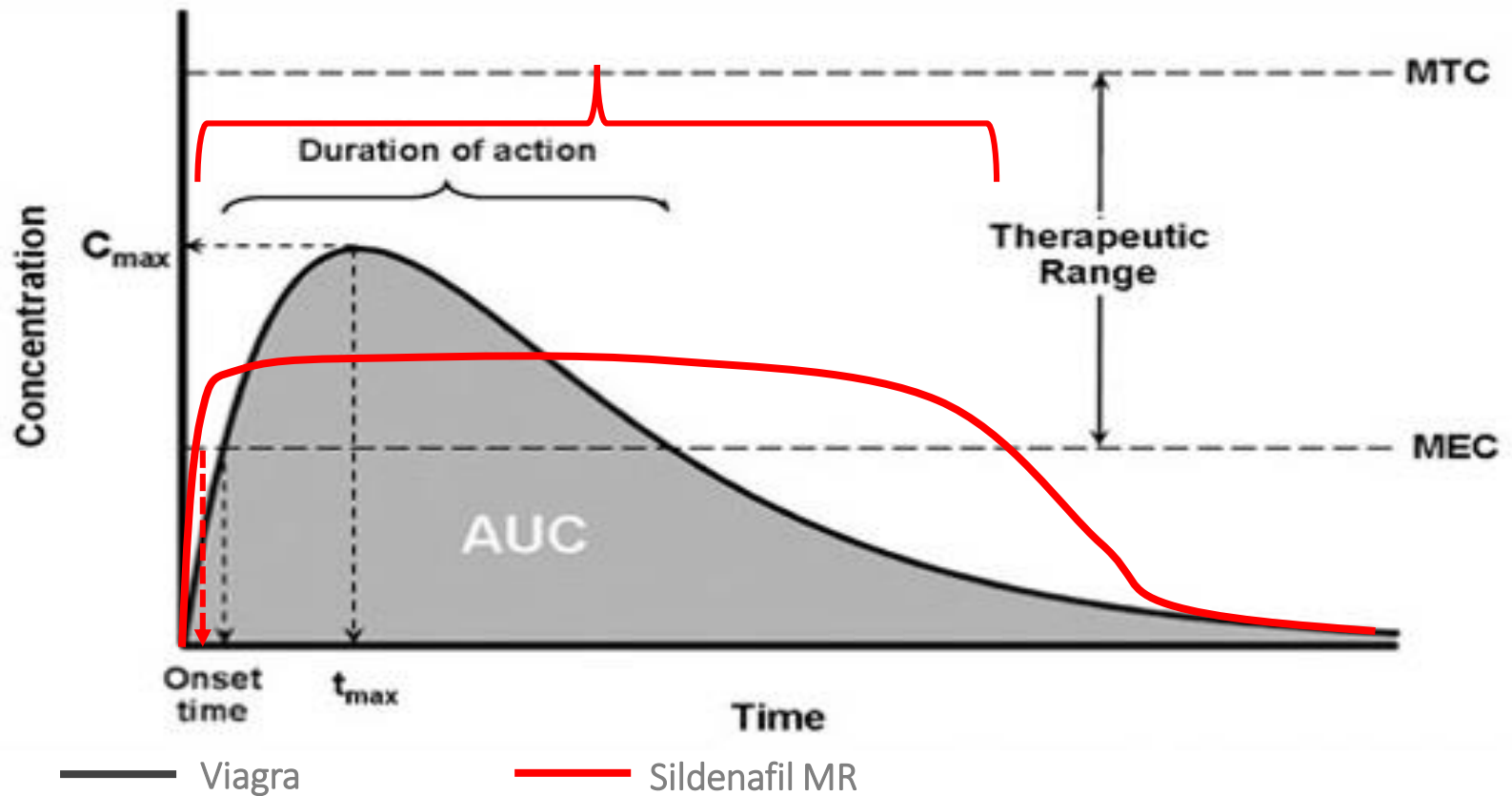
- Sildenafil MR is specifically developed to meet the needs of those patients suffering from ED

	Onset of action	Duration	Affected by food	Market size **
<b>Sildenafil MR (N4 Pharma)*</b>	<b>15-30 mins*</b>	<b>12-20 hours*</b>	<b>No</b>	<b>In development</b>
Viagra (Sildenafil)	1 hour	4-6 hours	Yes	\$1.6bn
Cialis (Tadalafil)	2-4 hours	36 hours	No	\$2.5bn
Levitra (Vardenafil )	40-50 mins	5-7 hours	Partial	\$0.2bn
Stendra (Avanafil)	15-20 mins	4-6 hours	Partial	n/a
MED2002 Gel (Futura)	5-10 mins	30 mins	No	In development

\* Target \*\* Source: Evaluate Pharma 2016

# Target Product Profile

- Sildenafil MR targets a flatter faster onset and longer lasting plasma concentration in the effective therapeutic window

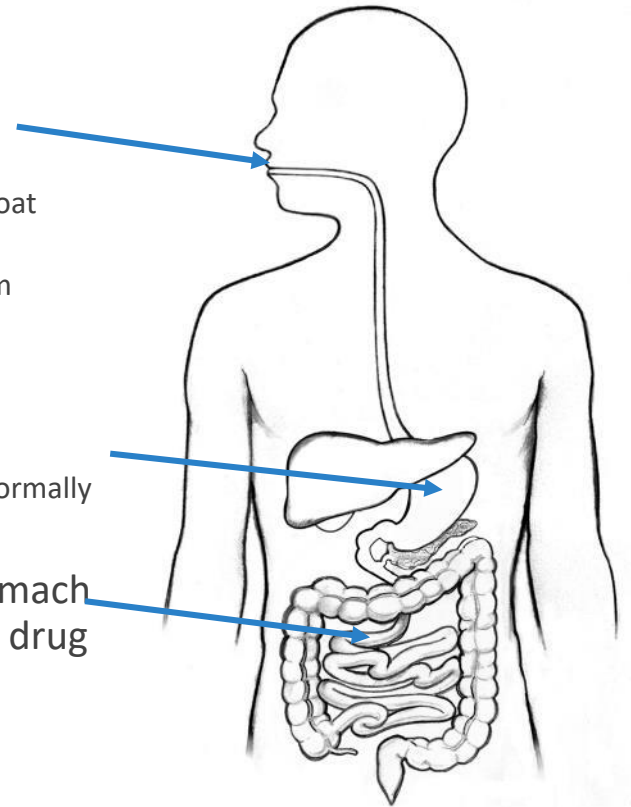


# Our sildenafil Product

- Sildenafil MR is a multiple action tablet comprised of an inner core of sildenafil citrate (80%) with an outer coat of sildenafil base (20%)



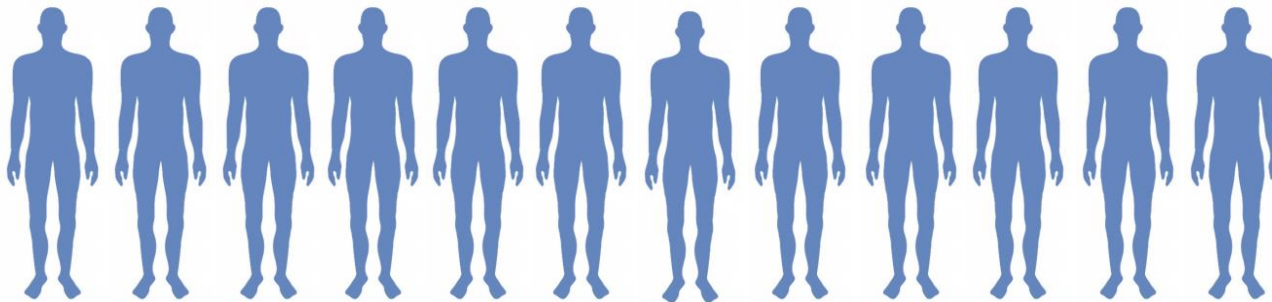
- Sildenafil MR placed under the tongue for 60 seconds
  - Initial release of sildenafil as outer coat partially dissolves
  - Delivers drug straight to bloodstream bypassing stomach
  - Rapid onset of action
- Tablet then swallowed
  - Remainder of outer coat dissolves normally in the stomach
- Inner core dissolves slowly in stomach and small intestine to enable the drug to last longer





# Commencement of human clinical trials

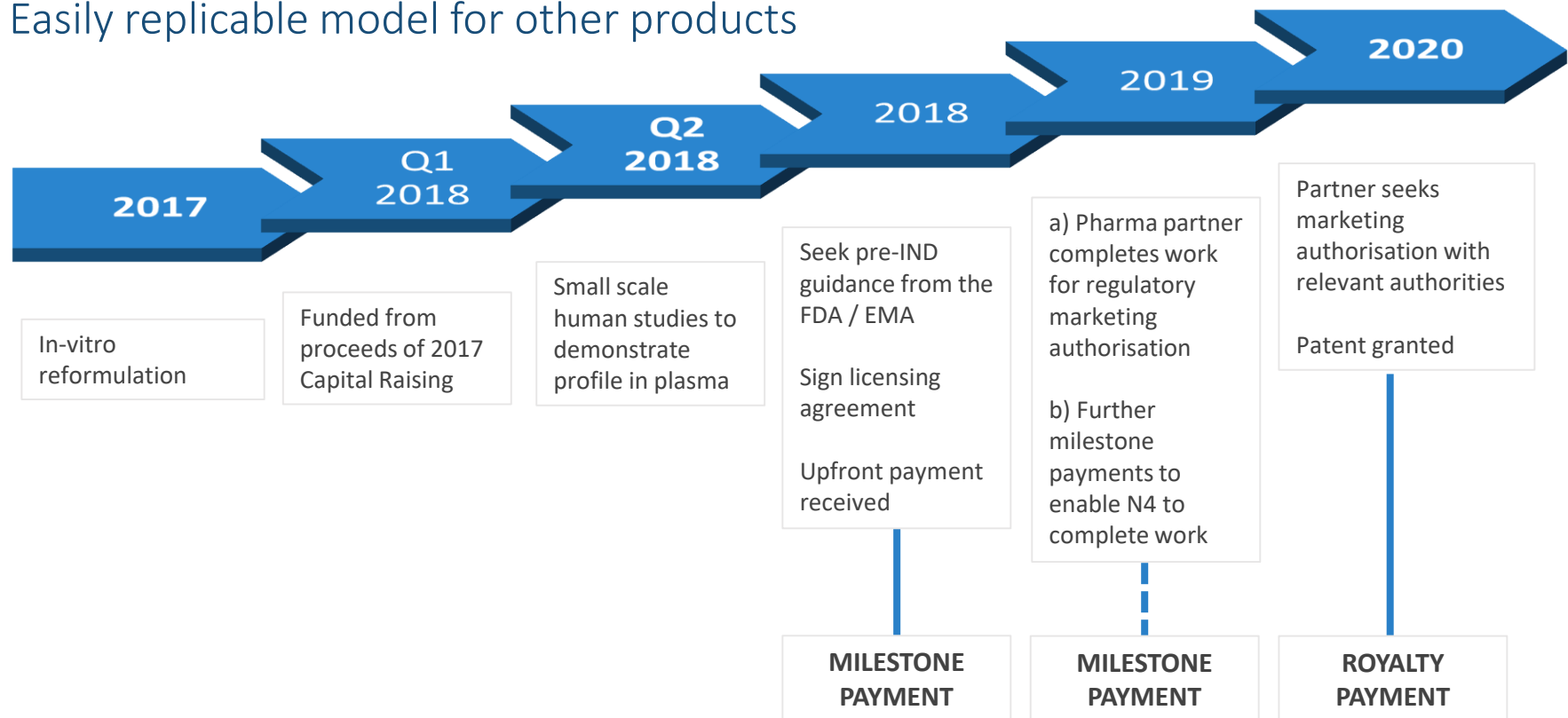
- Lead product sildenafil MR commenced proof of concept clinical trial – Q2 2018
- Four way crossover study in 12 healthy subjects
- Comparing reformulated 100mg sildenafil to Viagra 50mg in both fed and fasted conditions
- Aim to measure the level of the drug in plasma achieved by the reformulated product
- Data gathered establishes whether N4 Pharma's reformulation has been sufficiently successful




# Route to Market

## Commercialisation of sildenafil

Easily replicable model for other products

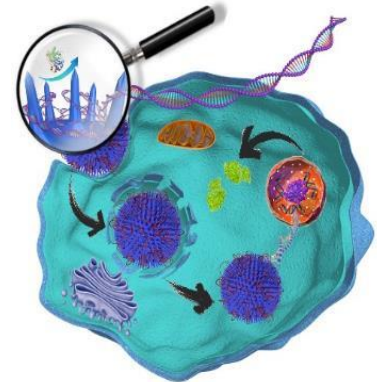




Nuvec<sup>®</sup>: Silica  
nanoparticle (SiNP)  
delivery system:  
For vaccines and  
therapeutics

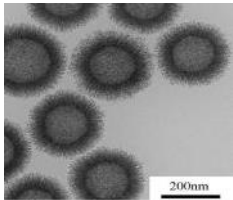
# What is Nuvec®?

- A novel nucleic acid delivery system for vaccines and biological therapeutics
- Mesoporous silica nanoparticles with unique structure designed as superior alternative to lipid systems and viral-like delivery vectors
- Addresses issues typically affecting current vectors:
  - High capacity loading of nucleic acids
  - Excellent transfection properties
  - No payload leakage, with evidence of protection from complete degradation
  - Excellent preclinical tolerability without major unwanted immune responses
    - crucially no tracking to the liver

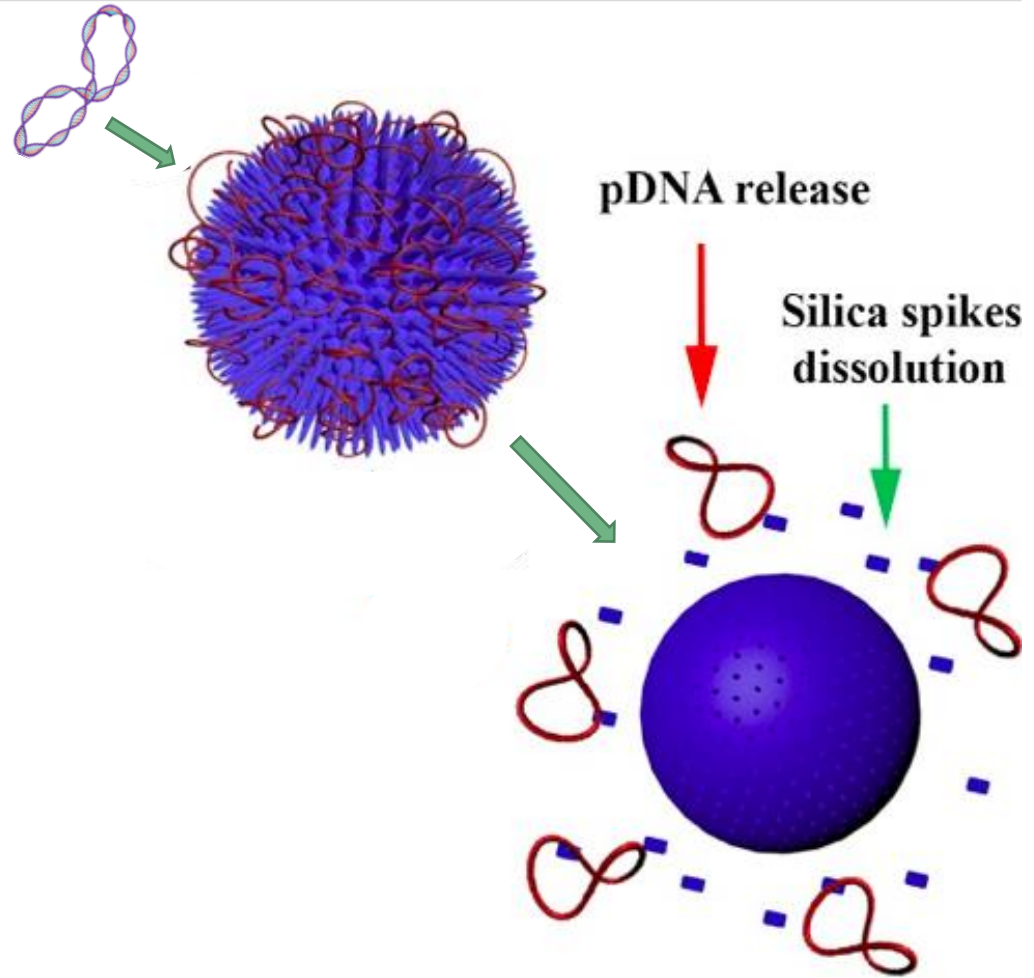


# Unique Design

- Hollow nano silica particles
- Ranging from 80nm to 500nm in diameter

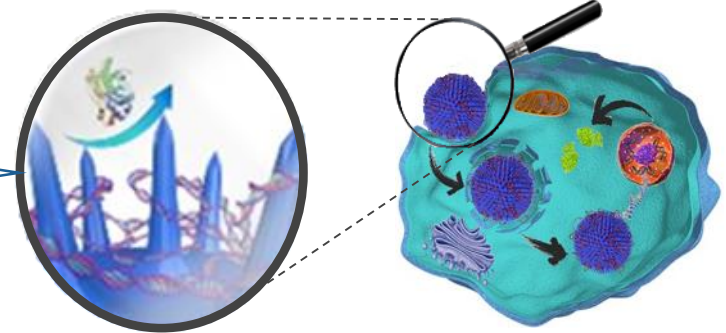


- DNA has a unique looped structure
- Difficult to attach to solid particles
- Nuvec®
  - Hollow silica sphere
  - Covered in thin silica spikes
  - Spikes trap the DNA
  - Spikes dissolve to release DNA
  - Particle then dissolves naturally in the body

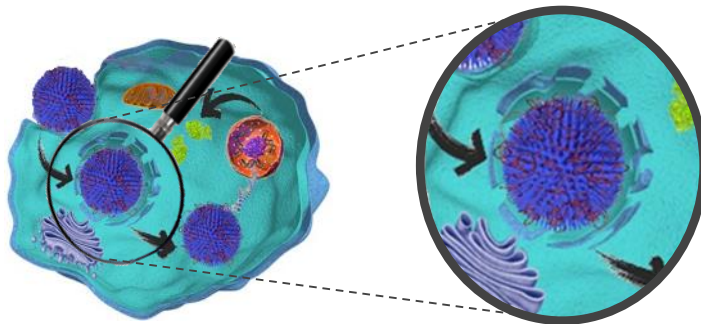


# Mechanism of action to deliver DNA/RNA

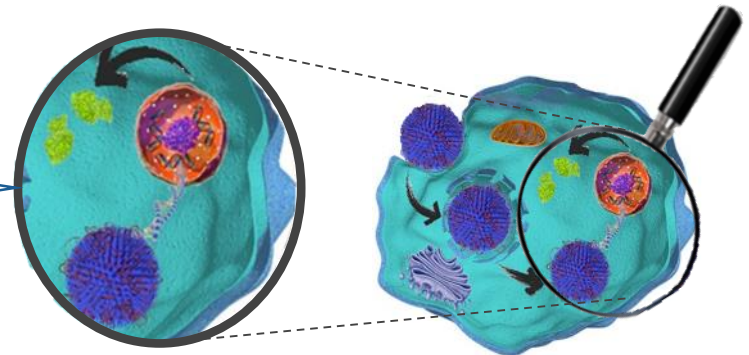
- Spikes trap and protect nucleic acid
- SiNPs attach to cell like Velcro and enter cell through process called pinocytosis



- Once inside SiNPs release contents into cytoplasm



- Nucleic acid sequence dissociates from SiNPs to allow generation of therapeutic proteins
- SiNPs degrade into harmless components



# Nuvec<sup>®</sup> Further learnings since RTO

- Shows great promise as a non viral delivery vector for DNA / RNA
  - Key advantages compared to lipid and viral systems
- High capacity loading and protection of nucleic acid for effective delivery
- Excellent localised *in-vivo* transfection efficiency
- No major systemic toxicity observed even with very high doses
  - SiNPs do not appear to induce high levels of inflammatory cytokines
- Internal studies currently focused on pDNA
  - mRNA studies planned for 2018 via collaborations
- Further research in progress
  - Dose response
  - Immune response
  - Stability
  - Toxicology tests



# Size of the Opportunity

Significant opportunities exist for a new novel delivery system

- The total market size for nanotechnology drug delivery was valued at \$79 billion in 2012 and is expected to reach a value of \$178 billion in 2019
- Large deal space:
  - Crescendo Biologics (private company)
  - Platform technology with multi-target application using special antibodies for cancer
  - Signed deal in 2016 with Takeda
    - \$36m upfront payment, investment and research funding, along with preclinical milestones
    - Potential for further revenue through milestones and royalties up to \$754m
- Nuvec research will inform exactly how we specialise in this market



# Route to Market

## Partnership opportunities with Nuvec®



- Nuvec® has wider potential for investigation other than just cancer vaccines
  - Cancer therapeutics
  - Immunotherapy
  - Reduction in Antibiotic resistance

# Summary

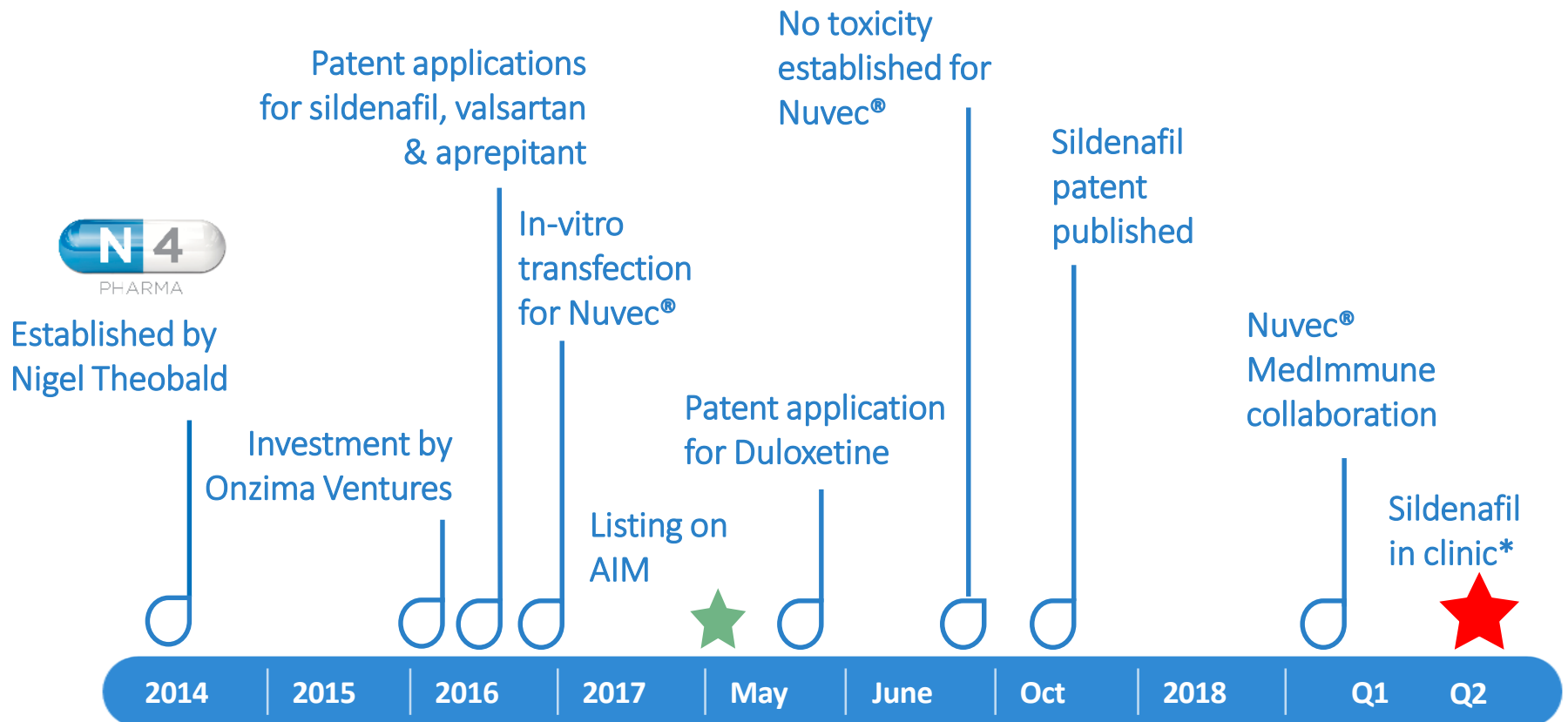
- Two divisions with similar business models
  - Targeting significant addressable markets
- Multiple collaboration opportunities
- Lead generic product, Sildenafil MR, in clinic
- Lower risk, less costly, quicker to market than traditional high risk drug development model
- Strong patent portfolio and IP
- Delivery system technology with huge potential for cancer vaccine and therapeutic market
- Highly experienced management team
  - Supported by experienced consultants with specialist knowledge in appropriate disciplines
- Early Nuvec® collaboration with MedImmune





# Appendix

# History of N4 Pharma



\*Expected results end Q2 2018

# Existing Pipeline and Strength of IP

## Slidenafil MR

Seeking to improve the speed at which sildenafil MR takes effect whilst extending the duration of action

- UK patent filed 02 April 2016
- PCT patent filed 31 March 2017
- PCT patent published -5 October 2017

## Nuvec®

Silica nanoparticle non-viral delivery system for DNA and RNA

- Patent licensed from University of Queensland
- PCT patent filed 20 October 2016
  - National phase applications October 2017

## Sartans

Used for the treatment of hypertension

- UK patent filed 01 July 2016
- PCT patent filed 03 July 2017

## Single Dose Hepatitis B

Sub unit vaccine – single dose reformulation of Hepatitis B surface antigen

Exclusive agreement with University of Queensland  
On hold- silica development focused on nuvec®

## Duloxetine

Reformulation of anti-depressant drug for the treatment of premature ejaculation.

- UK patent filed 11 May 2017

## Aprepitant

An anti-emetic drug used in oncology.

- UK patent filed 21 December 2016

# The Board

## **Nigel Theobald**

*Chief Executive Officer*

25 years' experience in healthcare & building businesses

Previously head of healthcare brands at Boots UK Limited

Grew Oxford Pharmascience Group to a £40m market cap company and managed the IPO on AIM

## **Paul Titley**

*Executive Director*

Over 40 years' experience in pharmaceutical industry

Built R5 Pharmaceuticals Limited into a profitable business leading to its acquisition by Aesica Pharmaceuticals, which itself was acquired by Consort Medical for £230 million in 2014

## **David Templeton**

*Non-Executive Chairman*

Experienced R&D manager with specific expertise in early clinical development

Previously worked for Pfizer, Xenova, Smithkline Beecham and GSK

Appointed as director of clinical pharmacology of Eisai Limited in 2007

## **Luke Cairns**

*Non-Executive Director*

16 years' experience in corporate finance and former head of corporate finance and MD at Northland Capital Partners

Founder of LSC Advisory Limited

Associate of the Chartered Institute of Secretaries