

VECTUS
BIOSYSTEMS
(ASX:VBS)

vectusbiosystems.com.au/

INVESTMENT HIGHLIGHTS



**3 FIRST IN CLASS
ASSETS ADDRESSING
MAJOR UNMET
THERAPEUTIC NEEDS**

VB0004 – Addressing Systolic Hypertension, cardiac, renal and pulmonary fibrosis, possible orphan indication for scleroderma

VB4-A32 – Addresses liver fibrosis, restored normal liver architecture in NASH/ASH models

VB4-A79 – Addresses pulmonary fibrosis from all causes except scleroderma where BP lowering probably required



**LEAD PROGRAM VB0004
CONTINUES POSITIVE
PROGRESS THROUGH
PHASE 1**

Completion of Single Ascending Dose (S.A.D) study with no adverse events observed

Completion of 14 day Multiple Ascending Dose (M.A.D) study with no significant adverse events

Interim data shows potential for single daily dose of VB0004 to treat various chronic conditions



**EXCEPTIONAL
PATENT PORTFOLIO
ENCOMPASSING A LIBRARY
OF > 1,000 COMPOUNDS**

VB0004 Patent granted in all major jurisdictions including USA, Europe, Japan, Peoples Republic of China, Republic of South Korea, Russian Federation, as well as Australia, Israel, Philippines, South Africa, Canada, ARIPO



**MANUFACTURING
SECURED ADDING
TO COMMERCIAL
POTENTIAL**

GMP manufacturing and scale up fully validated at two international centres

Broad pharmaceutical engagement established with both tier-1 and mid-size potential licensees

Targeting first-in-class reimbursement for large franchise indications



CORPORATE SNAPSHOT



KEY METRICS

ASX	VBS
Shares on issue	47.25m
Market Capitalisation	\$45.4m
Share Price (28/11/22)	\$0.96
52-week trading range	\$0.90 - \$1.70

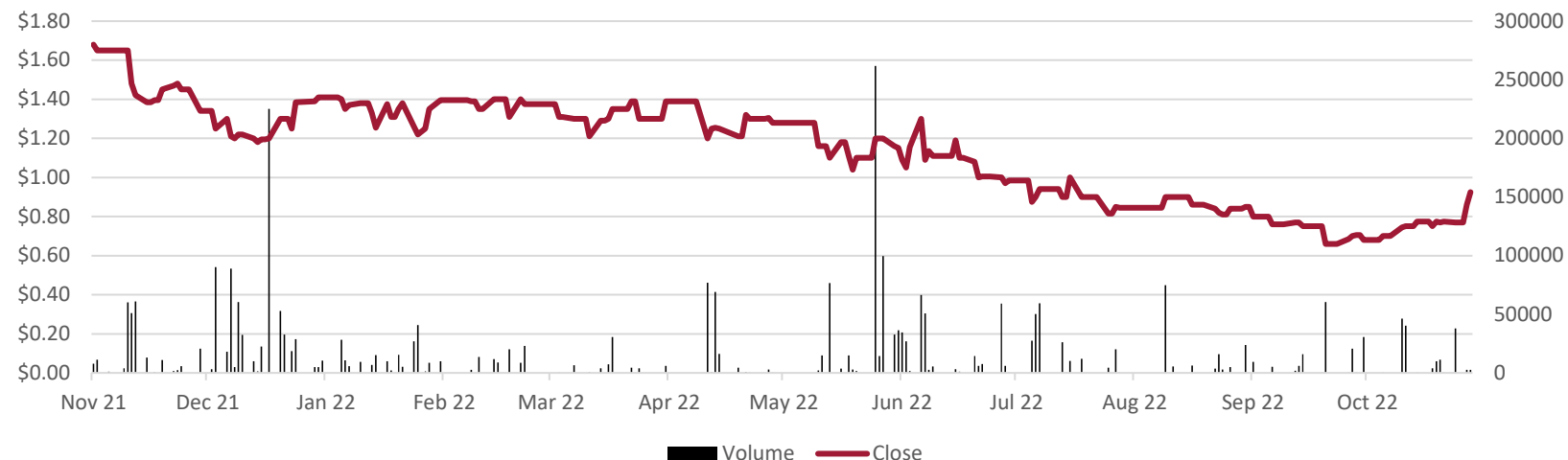
SHAREHOLDER BREAKDOWN

19.04%	Board of Directors
39.94%	Top 20
41.23%	Other

HIGHLY EXPERIENCED BOARD & MANAGEMENT

Dr Ronald Shnier Non-Executive Director and Chairman	Mr Maurie Stang Non-Executive Director and Deputy Chairman	Dr Karen Duggan Executive Director and Chief Executive Officer	Mr Peter Bush Non-Executive Director	Dr Susan Pond Non-Executive Director
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SHARE PRICE PERFORMANCE – LAST 12 MONTHS





Fibrosis is the replacement of functional tissue by scar tissue, usually as a result of injury, and is the pathology which underlies:



HEART FAILURE

(largest single item on US health care budget \$US32b in 2013)



LIVER FAILURE

(40% of population of China, India and South East Asia are affected)



KIDNEY FAILURE

(Dialysis and renal transplant costs in the US reached \$49.2b in 2011)



RESPIRATORY FAILURE

(pulmonary fibrosis)

FIBROSIS REVERSAL, A MAJOR UNMET NEED

Fibrotic disease
contributes to more than

40%

of all deaths worldwide

WHY VECTUS?



Vectus is strongly positioned with a range of transformational, orally doseable small molecules that target disease progression, aimed at meaningful functional changes & reversing existing damage to key cardiovascular organs

With continued success these agents would address the largest therapeutic indications, with no known direct competition & first-in-class reimbursement



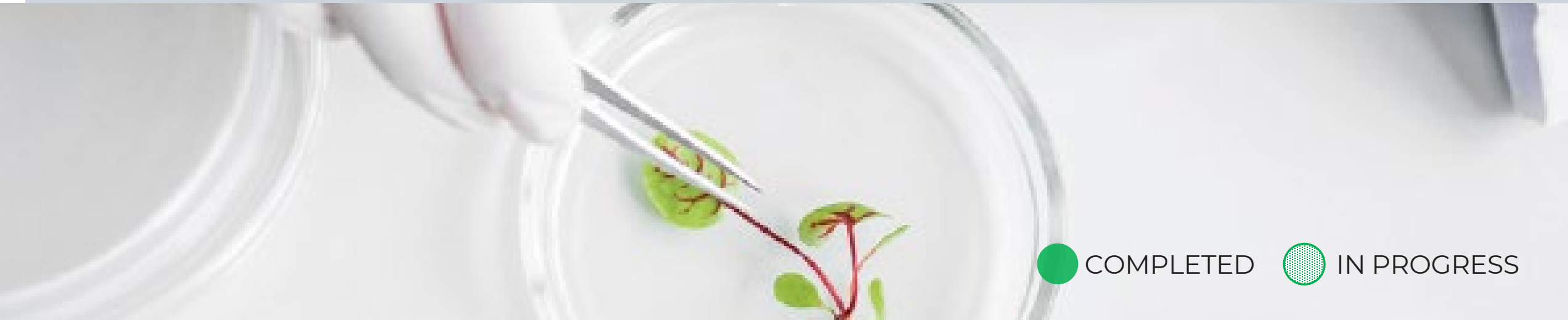
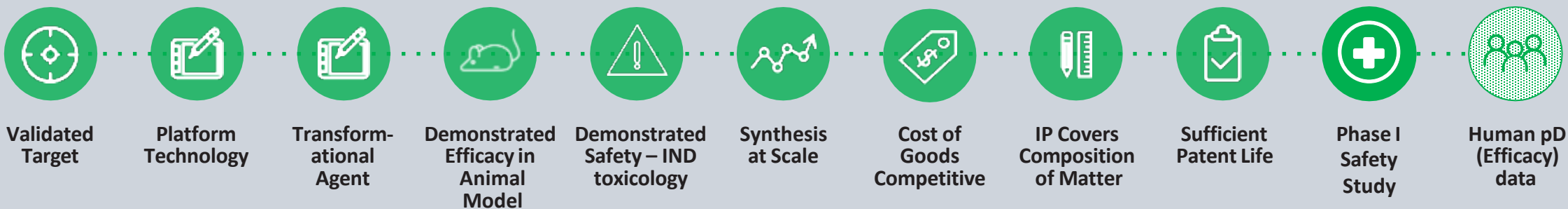
2023 CATALYSTS



VB0004	<ul style="list-style-type: none">• Completed Phase 1a Human Safety Trials• Phase 1B Human Trial in progress• Leverage engagement with global pharmaceutical companies
VB4-A32	Undertake GMP synthesis (Assymchem), with IND toxicology studies to follow
NEW EMERGING LEADS	Accelerate work on detailed mechanisms of action for VB4-A32, VB4-A79
EXPAND	Facilities and resources to undertake a broader drug development program
INVESTIGATE	Candidates from Vectus' extensive patented library for roles in other fibrotic / protein accumulative diseases such as osteoarthritis, retinal fibrosis, Alzheimer's disease
COMMERCIALISE	Broaden Accugen commercial roll out with Vectus' breakthrough qPCR platform



PHARMA LICENSING CRITERIA



 COMPLETED  IN PROGRESS



Treatment with VIP reversed cardiac fibrosis in multiple animal models data from one was published in the paper entitled “Vasoactive intestinal peptide reverses existing myocardial fibrosis in the rat”

European Journal of Pharmacology 862 (2019) 172629

Contents lists available at ScienceDirect

European Journal of Pharmacology

journal homepage: www.elsevier.com/locate/ejphar

Full length article

Vasoactive intestinal peptide infusion reverses existing myocardial fibrosis in the rat

Karen A. Duggan^a, George Hodge, Juchuan Chen, Tegan Hunter¹

^aVictus Biopharm, North Ryde, Australia

ARTICLE INFO

Keywords:
 Heart failure
 Myocardial fibrosis
 Vasoactive intestinal peptide

ABSTRACT

Congestive cardiac failure has become one of the major health challenges of the 21st century and new therapies are needed to address this problem. The concentration of vasoactive intestinal peptide (VIP) in the heart has been shown to decrease as fibrosis (the pathology leading to heart failure) increases and to become undetectable in end stage cardiomyopathy. We sought to determine whether replenishment of myocardial VIP might treat myocardial fibrosis and therefore represent a new therapeutic target.

Wistar Kyoto rats on a high (4.4%) salt diet were randomised to zero time control, 4 week infusion of VIP (5 pmol/kg/min) or vehicle control infusion. Myocardial VIP concentration was measured by radioimmunoassay, fibrosis was quantitated by computerised histomorphometry and changes in pro-fibrotic mediators were measured by quantitative q-PCR.

Myocardial VIP increased significantly in VIP treated rats compared with vehicle treated controls ($P < 0.01$) while fibrosis in the VIP treated rats was significantly lower than in both the zero time control ($P < 0.05$) and

Treatment with VIP was also found to reverse interstitial fibrosis in the kidney in multiple animal models data from one was published in the paper entitled “Vasoactive intestinal peptide infusion reverses existing renal interstitial fibrosis via a blood pressure independent mechanism in the rat”

European Journal of Pharmacology 873 (2020) 172879

Contents lists available at ScienceDirect

European Journal of Pharmacology

journal homepage: www.elsevier.com/locate/ejphar

Full length article

Vasoactive intestinal peptide infusion reverses existing renal interstitial fibrosis via a blood pressure independent mechanism in the rat

Karen A. Duggan^a, George Hodge, Juchuan Chen, Sofie Trajanovska¹, Tegan Hunter²

^aVictus Biopharm, North Ryde, Australia

ARTICLE INFO

Keywords:
 Renal failure
 Tubulointerstitial fibrosis
 Vasoactive intestinal peptide

ABSTRACT

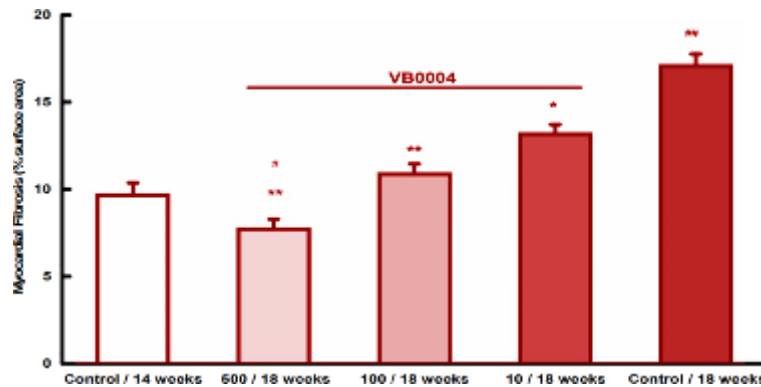
Dialysis requiring renal failure is a silent epidemic. Despite an annual mortality of 24% the dialysis population has increased by 1–4% per annum. Regardless of the initial injury, tubulointerstitial fibrosis is a feature of the renal pathology and it inevitably correlates with declining renal function. Current agents display little efficacy against tubulointerstitial fibrosis. Clearly, therapies effective against tubulointerstitial fibrosis and able to preserve kidney function are needed. Vasoactive intestinal peptide (VIP) has been shown to reverse pre-existing cardiac fibrosis. We sought to determine whether VIP is effective in tubulointerstitial fibrosis. Spontaneous hypertensive rats (SHR) on a 2.2% salt diet were randomised to zero time control, 4 week infusion of VIP (5 pmol/kg/min) or vehicle control infusion. A fourth group, to match the blood pressure reduction achieved in the VIP infused group was included. Fibrosis was quantitated by computerised histomorphometry, changes in pro-fibrotic mediators were measured by quantitative q-PCR and macrophage activation assessed by cyclic adenosine monophosphate (c-AMP) response element luciferase with VIP. Tubulointerstitial fibrosis in the VIP treated



VB0004 has been shown to:

- Rescue cardiac tissue damaged by fibrosis
- Repair existing cardiac damage
- i.e. VB0004 is transformational

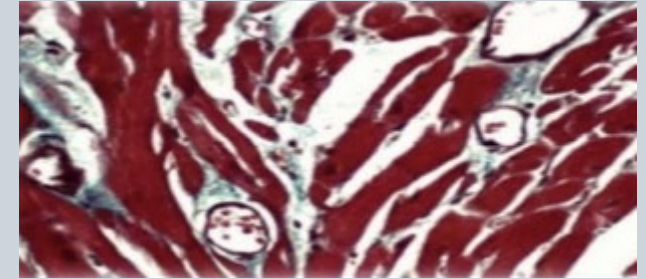
Treatment with VB0004 at 3 Doses



At the highest dose (500pmol/kg/min), VB0004 reversed pre-existing fibrosis, while a dose response effect on the level of fibrosis is apparent

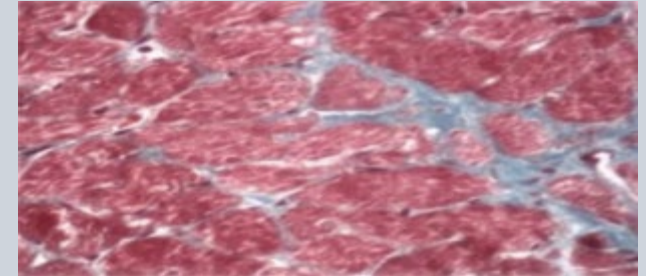
14-Week Control

Fibrous tissue (blue staining) is visible around blood vessels and extending between muscle fibres



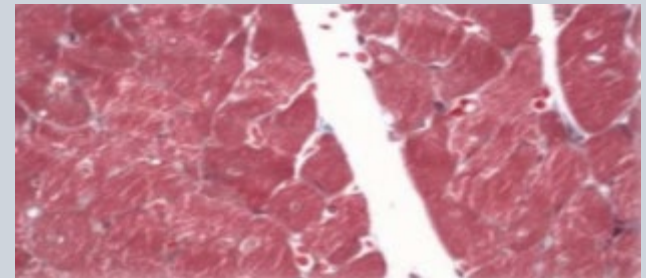
5% Ethanol 18-Week Control (Vehicle Control For VB0004)

Fibrosis visible as blue stained tissue is present throughout the section



Heart At 18 Weeks After 4-week Treatment with VB0004 (500 Pmol/Kg/Min)

Minimal fibrosis is visible; normal architecture has been restored

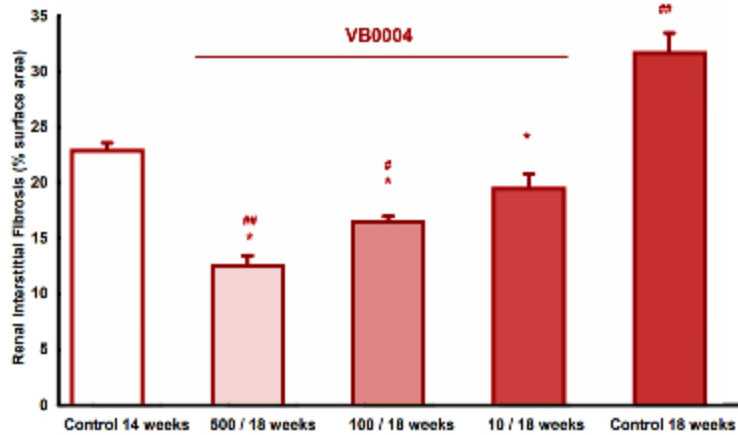




In the kidney VB0004 has been shown to:

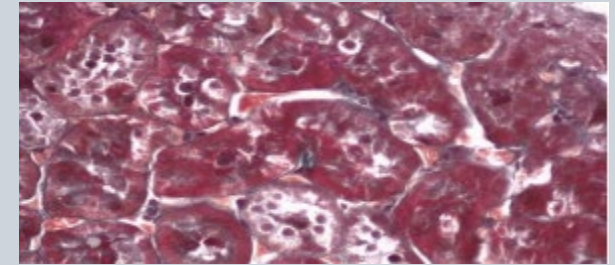
- Reverse renal interstitial fibrosis at all doses
- Restore normal architecture at all doses (i.e. VB0004 is considered transformational)

Treatment with VB0004 at 3 Doses



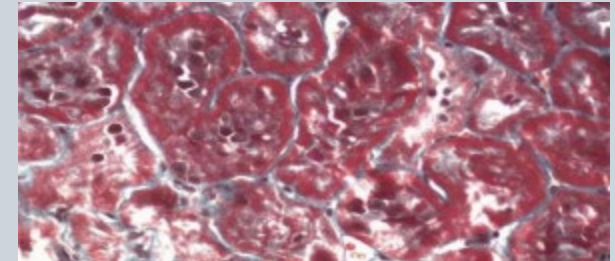
14-Week Control

Fibrosis (blue) partially surrounds some but not all tubules



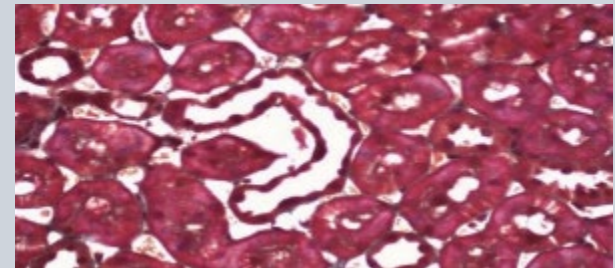
5% Ethanol 18-Week Control (Vehicle Control For VB0004)

Fibrosis has progressed to surround most tubules



Kidney At 18 Weeks After 4-week Treatment with VB0004 (500 Pmol/Kg/Min)

No fibrosis visible

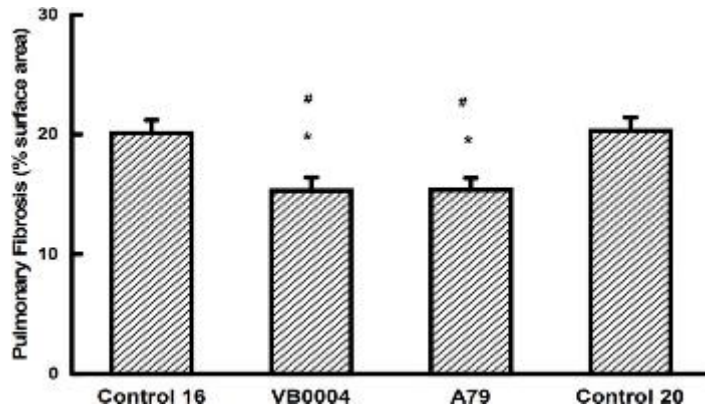




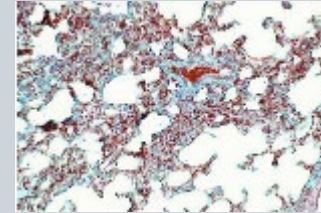
In the lung:

- VB0004 reversed fibrosis present 2 weeks after treatment with bleomycin (an anticancer drug)
(i.e. VB0004 also transformational in the lung)

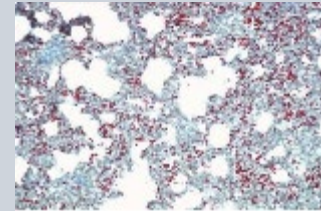
Treatment with VB0004 and VB4-A79



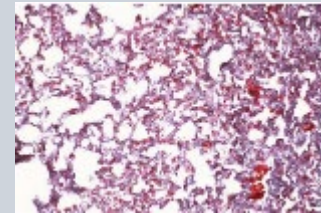
16-Week Control



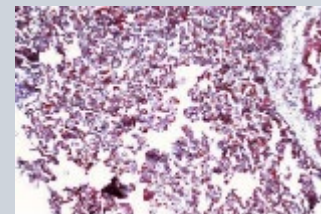
20-Week Control



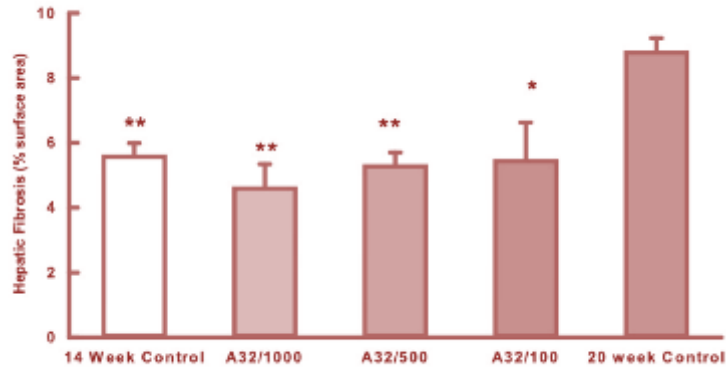
VB4-A79 at 20 weeks



VB0004 at 20 weeks

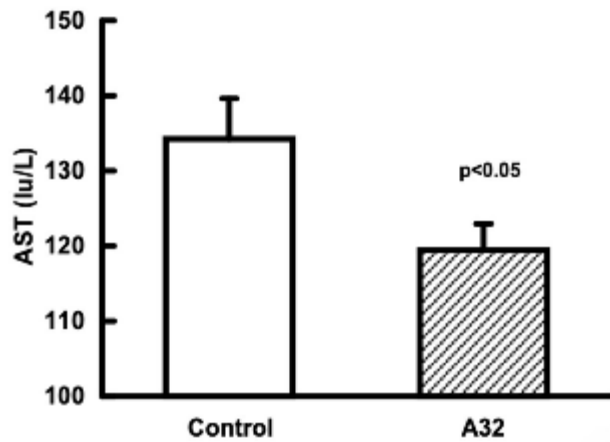


VB4-A32 & HEPATIC CIRRHOSIS

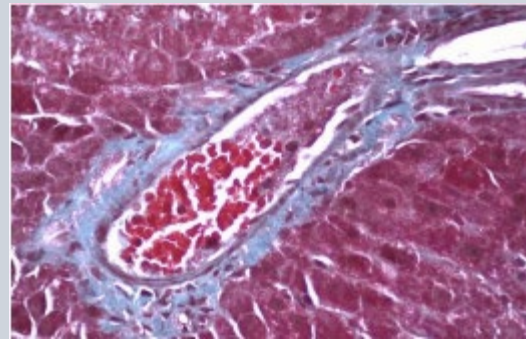


VB4-A32 demonstrated ability to:

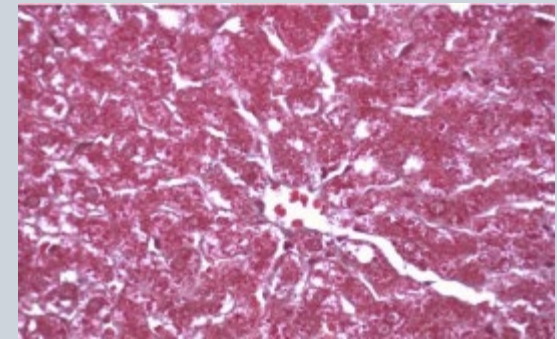
- Reduce peri-portal fibrosis in the liver in a dose dependent manner (above left)
- Improve liver function tests (below left)



20-Week Control



A32 20 Weeks





01

FIRST GMP SYNTHESIS BY GLYCOSYN

- Yield increased as scale increased
- VB0004 manufactured to 5kg scale
- Cost efficient at 5kg scale
< \$(US)0.05 per mg
- Estimated dose 1-5mg
- Stability studies – stable at 2 yrs (long shelf-life)



02

SECOND GMP SYNTHESIS BY ASSYCHEM

- Campaign planned to provide 3 validation batches
- Confirm consistency of the synthesis process
- Samples of all 3 will undergo 2 yr stability testing
- Meets FDA requirements for GMP manufacture for Phase 1 and 2 clinical trials



PHASE 1



Trial design – conventional Single Ascending Dose (SAD) & Multiple Ascending Dose (MAD)

Includes pharmacokinetic and pharmacodynamic studies

Expected outcomes – maximum tolerated dose, dose limiting toxicity (if present), pharmacokinetic (PK) and pharmacodynamic (PD) data

Syneos Health (Nasdaq SYNH) retained to write Investigator Brochure (IB), trial protocol and monitor Phase 1 trial

SAD completed with no significant adverse events, max tolerated dose 300mg

Healthy subjects 14 day MAD completed, no significant adverse events, max tolerated dose 100mg daily

PK – Tmax at 6-8hrs, half life 10-15hrs, no accumulation over 14 days

Affected individuals 2 groups 28 days 2 doses biomarkers identified - recruiting





Fist in class therapeutic

- VIP agonist

Transformational agent

- reverses existing fibrotic disease
- effective in multiple organs

Synthesis

- 3 steps
- cost competitive (\$0.05 /mg)

Stability

exceeds 2 years

Efficacy

- significantly greater reductions in SBP, cardiac and renal fibrosis at lower doses than current agents.

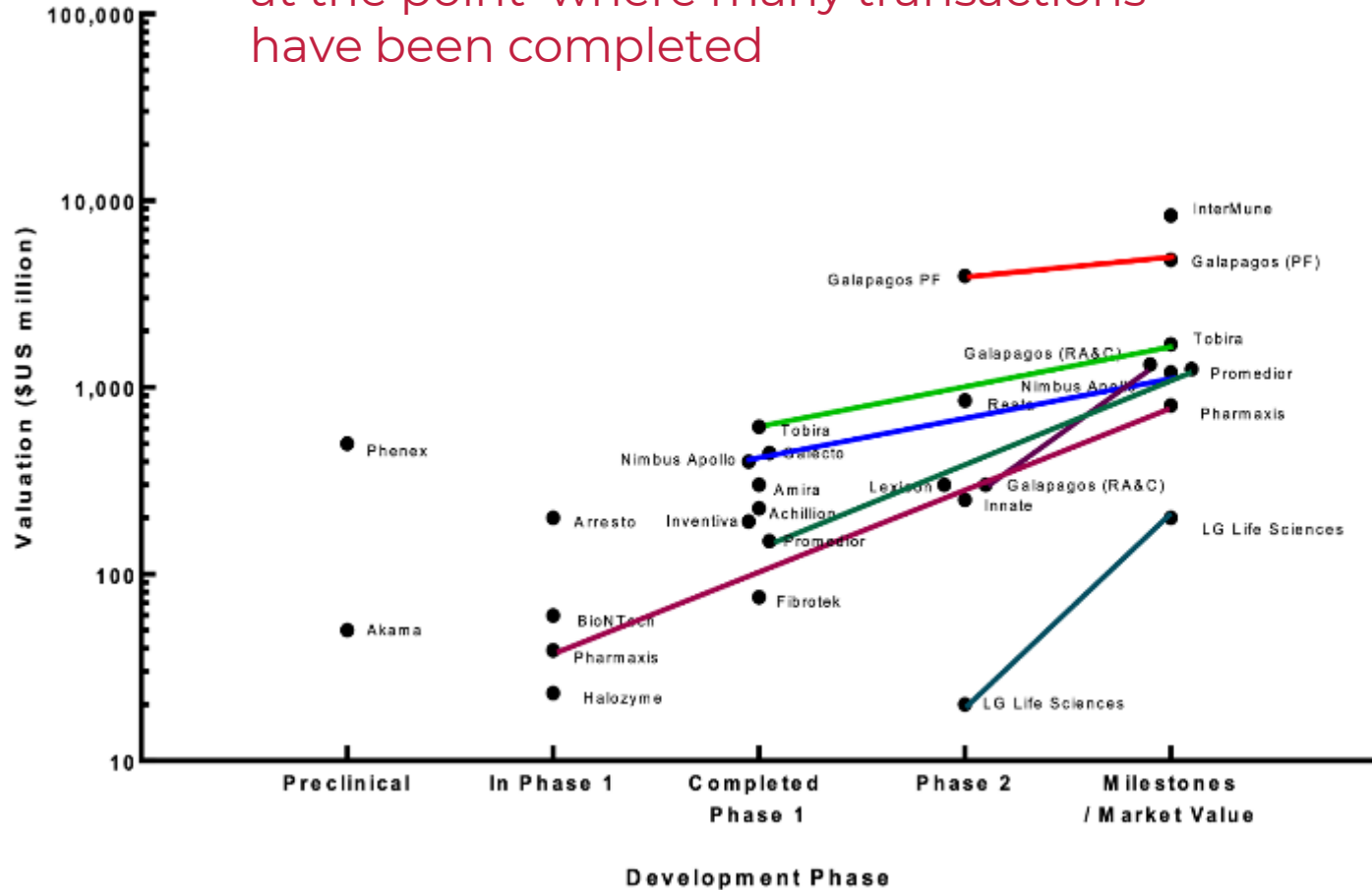
Potential role

- given significantly greater efficacy at lower doses than current agents may replace RAS blockers as the foundational agent for therapeutic regimens in cardiovascular and renal disease

COMPARABLE TRANSACTIONS



Successful Phase 1/1b would place Vectus at the point where many transactions have been completed

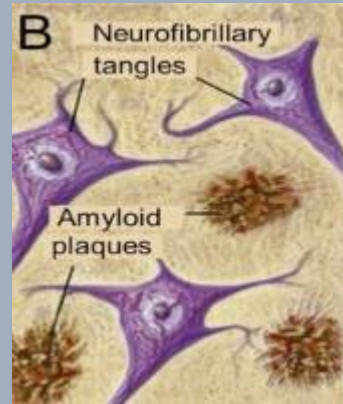
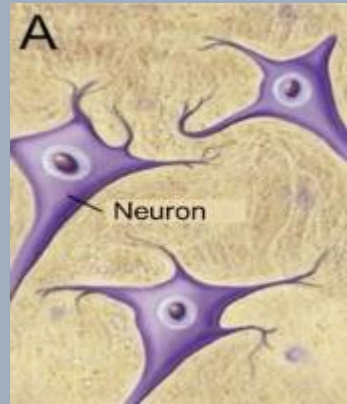


NEW INDICATIONS ALZHEIMERS & LONG COVID



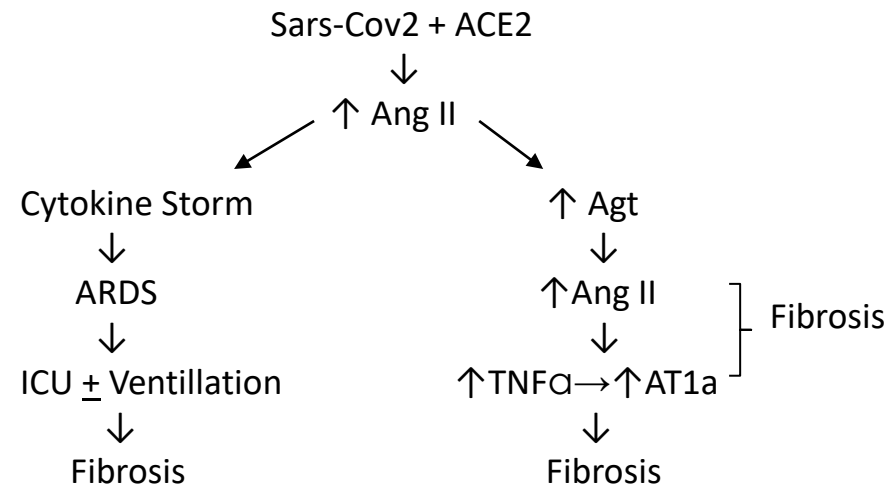
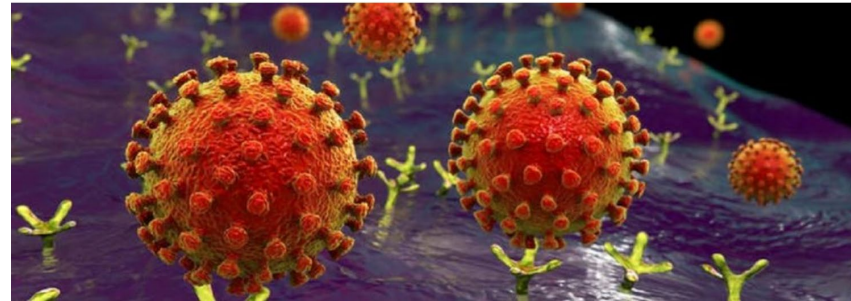
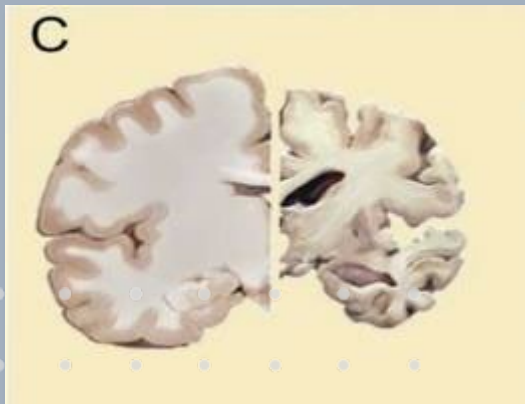
NORMAL

ALZHEIMER'S DISEASE

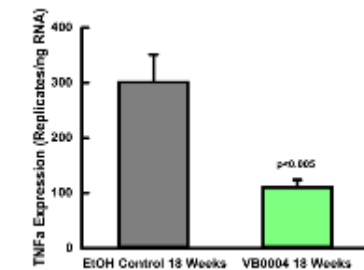
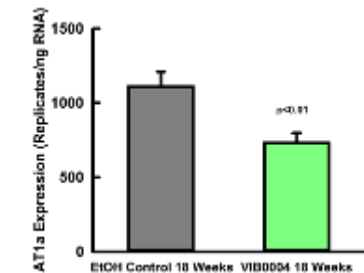
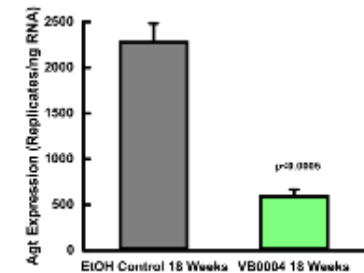


HEALTHY BRAIN

SEVERE ALZHEIMER'S



Ang II = Angiotensin II
 ARDS = Acute Respiratory Distress Syndrome
 Agt = Angiotensinogen, the Ang II precursor
 TNFα = Tumour Necrosis Factor alpha



PATENT PORFOLIO – LONG PATENT LIFE



VIP patents for heart, kidney and aortic fibrosis

— granted all jurisdictions

VIP fragment patents compositions and methods of use for hypertension, cardiac, renal and aortic fibrosis

— granted most jurisdictions

VB0004 compositions and methods of use for hypertension, cardiac and renal fibrosis

— granted Russian Federation, Israel, Singapore, ARIPO, Canada, Philippines, South Africa, Ukraine, Vietnam, Nigeria, Mexico, accepted in Indonesia

VB0004 library of approx. 70 related compounds compositions and methods of use for treatment of hypertension, cardiac and renal fibrosis

— granted US, Australia, China, Europe, Japan, Korea, Russia, Ukraine, Hong Kong, Vietnam, Singapore, accepted in South Africa, ARIPO, Brazil, accepted Mexico

VB4-A32 and library of related compounds compositions and methods of use for treatment of hepatic, cardiac and renal fibrosis

— granted US, Europe, Australia, South Africa

VB4-P5 and library of related compounds compositions and methods of use for treatment of renal cell death, renal fibrosis and hepatic fibrosis

— granted US, China, Australia, South Africa, accepted Europe, Japan, Russia, Israel

GMP method of synthesis VB0004

— granted USA, Australia, India, accepted Europe, China

VB4-A79 and related compounds compositions and use for treatment of pulmonary fibrosis

— granted Australia, China, accepted USA, Europe, Mexico

VB0001 and related compounds compositions and use for management of hypertension and fibrotic disease

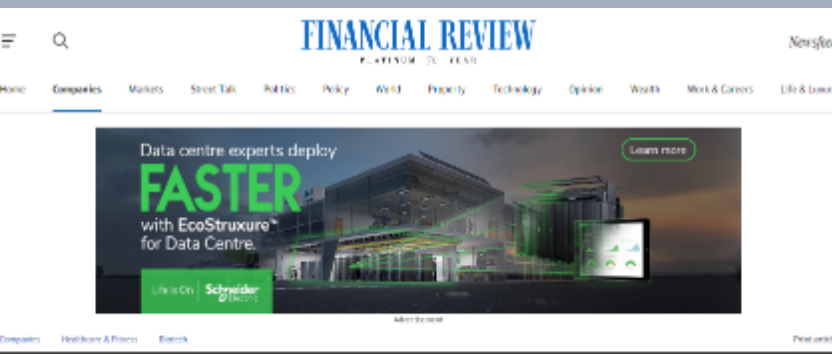
— PCT application

VB0002, VB0003 and VB0005 and related compounds compositions and use for management of hypertension and fibrotic disease

— national phase



IN THE MEDIA – VECTUS INCREASING MARKET RECOGNITION



Why medtech veteran Maurie Stang says the time is right for Vectus



Yolanda Hedberg
Reporter

May 25, 2022 - 4:20pm

Almost 15 years after the company was founded, medical technology veteran Maurie Stang is convinced 2022 is the tipping point for drug discovery company Vectus Biosystems.

The ASX-listed small cap, which surged more than 8 per cent on Wednesday, is developing a series of oral drugs based on a peptide in the intestine to treat cardiovascular, pulmonary and liver fibrosis, as well as systolic blood pressure.



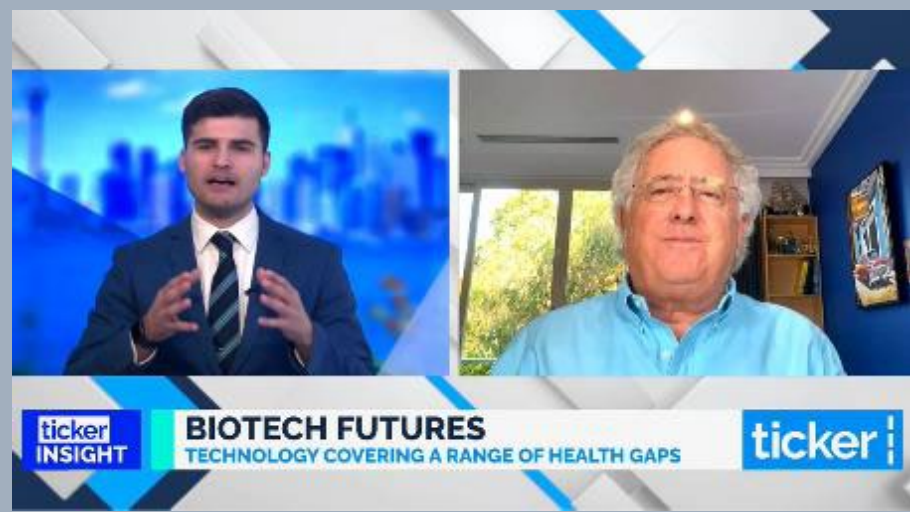
Dr Boreham's Crucible: Is Vectus Biosystems about to become a VIP for takeover suitors?

October 3, 2022 | Tim Boreham

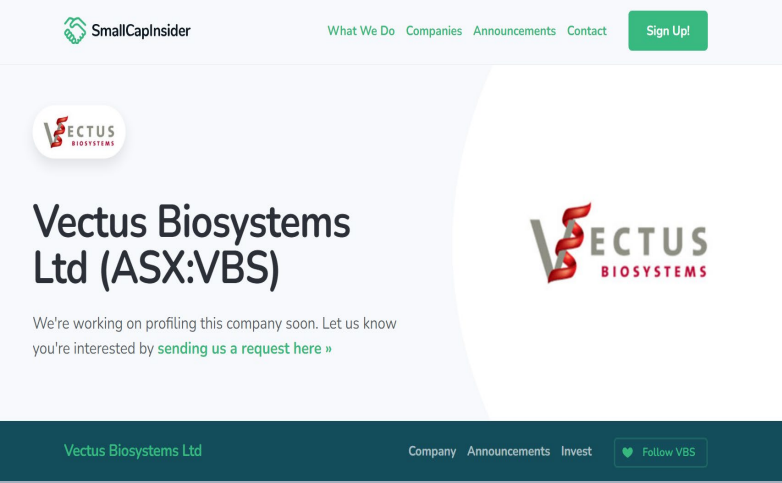


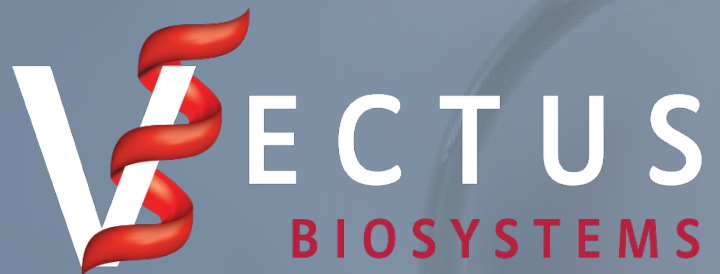
ASX Health Stocks: Vectus jumps 13pc on promising Phase 1 study results

May 25, 2022 | Eddy Sunarto



Vectus Biosystems progresses its human trials with lead cardiovascular drug candidate VB0004





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