



Investor Presentation | July 2023 Gary Phillips CEO

## **Forward looking statement**

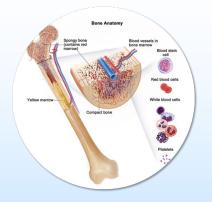
This document contains forward-looking statements, including statements concerning Pharmaxis' future financial position, plans, and the potential of its products and product candidates, which are based on information and assumptions available to Pharmaxis as of the date of this document. Actual results, performance or achievements could be significantly different from those expressed in, or implied by, these forward-looking statements. All statements, other than statements of historical facts, are forward-looking statements.

These forward-looking statements are not guarantees or predictions of future results, levels of performance, and involve known and unknown risks, uncertainties and other factors, many of which are beyond our control, and which may cause actual results to differ materially from those expressed in the statements contained in this document. For example, despite our efforts there is no certainty that we will be successful in developing or partnering any of the products in our pipeline on commercially acceptable terms, in a timely fashion or at all. Except as required by law we undertake no obligation to update these forward-looking statements as a result of new information, future events or otherwise.

# **Executive Summary**

- Pharmaxis is a clinical stage drug development company targeting inflammation, fibrosis and selected cancer indications with first in class or best in class small molecule drugs in markets of high value
- Global leader in fibrosis driven by lysyl oxidase enzymes having invested in a multi year research program leveraged with extensive external scientific collaborations
- Breakthrough data and supportive feedback from FDA provides clear pathway to commercial value in \$1bn myelofibrosis market
- Cash position at 31 March 2023 of A\$15m, plus 2023 R&D tax credit similar to 2022 (\$5m).

### FIRST IN CLASS ANTI-FIBROTIC DRUGS



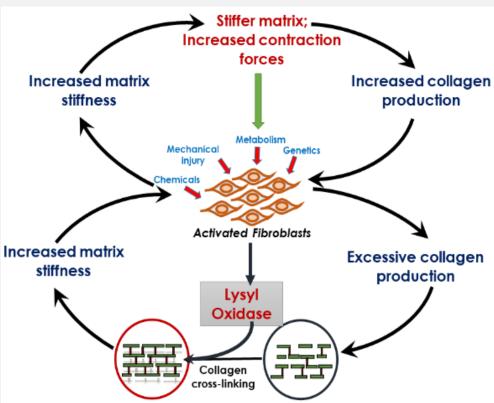


Clinical proof of concept that LOX inhibition reduces fibrosis achieved in two diseases in 2023

## Pharmaxis is the global leader in lysyl oxidase chemistry and biology

Multi year research program leveraged with extensive scientific collaborations worldwide has delivered 2 drugs in the clinic

### Lysyl oxidases are the final stage in fibrosis



Tissue stiffening due to increases in collagen and number of crosslinks which is a hallmark of fibrosis, is preventable through lysyl oxidase inhibition; at the heart of a true anti-fibrotic therapy

#### PXS-5505

- Oral dosage form four capsules twice a day
- Patent filed priority date 2018
- Strong pre clinical evidence in models of fibrosis and cancer
- INDs approved for myelofibrosis and hepatocellular carcinoma
- Potential in multiple cancer indications
- Phase 1 data demonstrates a safe, well tolerated drug that gives >90% inhibition of LOX enzymes

#### PXS-6302

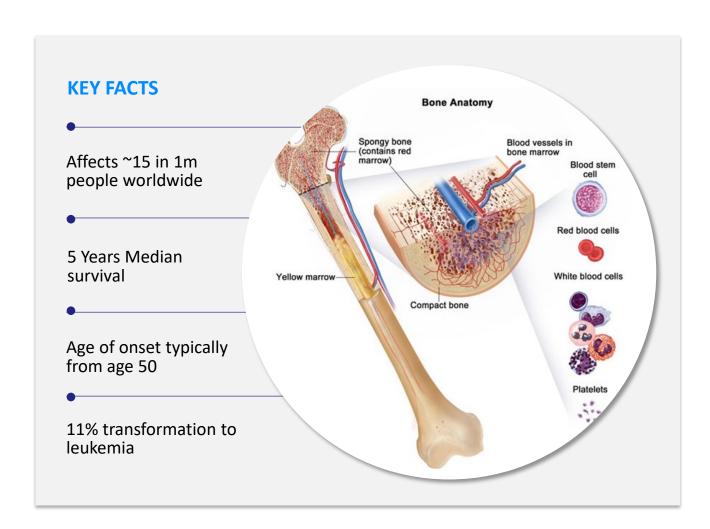
- Topical dosage form
- Patent filed priority date 2019
- Strong pre clinical evidence in models of skin fibrosis and scarring
- Potential in prevention of scar formation and modification of existing scars
- Phase 1a (healthy volunteer) data demonstrates a safe, well tolerated drug that gives full inhibition of LOX enzymes in the skin with minimal systemic exposure

# **Program Update**



# **Myelofibrosis**

A rare type of bone marrow cancer that disrupts the body's normal production of blood cells



**Primary Myelofibrosis** is caused by a build up of scar tissue (fibrosis) in bone marrow reducing the production of blood cells:

- Reduced red blood cells can cause extreme tiredness (fatigue) or shortness of breath
- Reduced white blood cells can lead to an increased number of infections
- Reduced platelets can promote bleeding and/or bruising
- Spleen increases blood cell production and becomes enlarged
- Other common symptoms include fever, night sweats, and bone pain

#### **Current Standard of Care; JAK inhibition**

- Symptomatic relief plus some limited survival improvement. 75% discontinuation at 5 years
- Median overall survival is 14 16 months after discontinuation

### **Commercial Opportunity**

 Current standard of care ; revenue ~US\$1b per annum

## **Myelofibrosis - PXS-5505 Phase 1/2a Trial**

6 month monotherapy study with meaningful safety and efficacy endpoints

**DESIGN** TREATMENT COHORT **ENDPOINTS** Phase 2a open label **Cohort expansion: Primary:** Safety and tolerability study to evaluate safety, PXS-5505 PK/PD, and efficacy (n = 24 subjects) 26 weeks **Secondary:** PK/PD JAK-inhibitor unsuitable\* Bone Marrow Fibrosis Grade primary MF or post-ET/PV **IWG** Response MF patients with: Spleen Volume Response • INT-2 or High risk MF Haematology requiring therapy Symptom score Symptomatic • BMF Grade 2 or greater

FDA granted orphan drug designation July 2020 and IND approved August 2020

20 sites across 4 countries (Australia, South Korea, Taiwan, USA)

Study recruitment commenced Q4 2021



# Myelofibrosis - PXS-5505 Phase 2a Trial (FINAL INTERIM DATA)

Very well tolerated with encouraging signs of clinical efficacy in JAK inhibitor unsuitable patients

### Study status

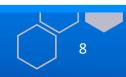
- 21 out of a targeted 24 patients have been enrolled
- 10 patients having completed 24 weeks of treatment

### Safety

- PXS-5505 has been well tolerated with no serious treatment related adverse events reported
- Majority of AEs were mild and not related to treatment
- 10 patients have dropped out of the study; none were treatment related

### Efficacy

- 5/9 evaluable patients\* had improved bone marrow fibrosis scores of ≥1 grade with 4 out of 5 fibrosis responders demonstrating stable hematological parameters and 3 out of 5 patients reporting symptomatic improvement
- 4 had an improvement in symptom score of >20%
- 7 had stable/improved hemoglobin (Hb) counts
- 8 had stable/improved platelet counts; 3 of these 8 patients entered the study with Grade 4 (potentially life-threatening) thrombocytopenia
- No spleen volume response (SVR35) was identified



# PXS-5505 Phase 2 Trial (MF-101); Expert review of interim data

### **Key Opinion Leader Review**

- "PXS-5505 continues to show not only an excellent safety profile but also promising clinical activity. The effect on bone marrow fibrosis is particularly exciting for a disease like myelofibrosis, where despite numerous years of research, we do not have any effective anti-fibrotic drugs."
- "It is encouraging to see that majority of 10 patients who completed 24 weeks of therapy also had improvements of symptoms and more importantly, stable or improved blood counts; including in those patients with severe thrombocytopenia."
- "These results support plans to continue clinical investigation of the agent, including combinations with JAK inhibitors where the lack of overlapping hematological toxicity would make PXS-5505 an ideal addon candidate."



Dr. Lucia Masarova
Assistant Professor, Department of
Leukemia at MD Anderson Cancer Center,
Houston

# PXS-5505 myelofibrosis clinical development plan: Regulatory update

#### FDA feedback:

- FDA Type C Meeting held in Q2 2023
- FDA reviewed all safety and efficacy data available at that time.
- Subject to protocol review FDA supported progression into a study in combination with a JAK inhibitor
- FDA provided guidance on the number of patients, treatment dosage, study duration and endpoints
- Trial protocol proposed to FDA
  - Uses existing trial sites; fast start up and minimal initiation costs
  - No dose escalation step; fastest route to meaningful data
- FDA feedback expected July 2023

## Hypertrophic and keloid scarring

Cutaneous scarring following skin trauma or a wound is a major cause of morbidity and disfigurement

#### **KEY FACTS**

•

100m patients develop scars in the developed world alone each year as a result of elective operations and operations after trauma

Hypertrophic scars and keloids are fibroproliferative disorders that may arise after any deep cutaneous injury caused by trauma, burns, surgery, etc.

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Hypertrophic scars and keloids are cosmetically and functionally problematic significantly affecting patients' quality of life



"In (preclinical) models of scarring we found that topical application of PXS-6302 reduces collagen deposition and crosslinking and improves scar appearance without reducing tissue strength. This is a unique way of modulating a critical stage in scar formation and maintenance and holds out great promise for the treatment of scars."

- Dr Mark Fear, UWA

 Mechanisms underlying scar formation are not well established; prophylactic and treatment strategies remain unsatisfactory

#### Current standard of care includes:

- Corticosteroids
- Surgical revision
- Cryotherapy
- Laser therapy
- 5-fluorouracil



#### Pre clinical evidence

 Treatment with PXS-6302 monotherapy demonstrates cosmetic and functional improvements to scarring in pre clinical models<sup>1</sup>

#### Clinical evidence

 3 month phase 1c in established scars demonstrates good tolerability, full inhibition of LOX in skin and marked change in scar composition

#### Commercial Opportunity

 Total scar treatment market in 2019 exceeded US\$19b. Keloid and hypertrophic scar segment ~US\$3.5b

### Established Scarring - PXS-6302 Phase 1c Trial (Solaria 2)

3 month monotherapy study to assess dosage, tolerability and efficacy endpoints

#### **DESIGN** PATIENT DEMOGRAPHICS **ENDPOINTS** Phase 1c 42 Adult patients (18-60) with **Primary:** an established scar > 1year: Safety and tolerability 3 month Average age of scar; 12.8 **Secondary:** • Objectives: **years** Characterize PK/PD\* Low to moderate severity parameters Confirm PK/PD\*, safety and efficacy of dose Included all surgery types. **Exploratory:** selected in dose escalation • Scar > 10cm<sup>2</sup>. Physical and visual skin and scar assessments Excluded patients with **Double blind placebo** acute skin conditions or controlled history of keloids

Investigator initiated study (sponsor UWA) - long term collaboration with UWA to research and develop PXS-6302 supported by Australian NHMRC grants

Single site study in Perth Australia

Study Completed March 2023

Study reported May 2023



### PXS-6302 Phase 1c Trial (Solaria 2); Top line results

- PXS-6302 was very well tolerated and demonstrated a good safety profile.
  - No serious adverse events were reported
  - Two patients withdrew from the study; reversible rash
- Mean inhibition of LOX activity 66% compared to baseline and placebo (p<0.001)
  - LOX measured 2 days post final dose
  - LOX is responsible for the cross linking of collagen fibres implicated in adverse scarring.
- Meaningful changes in the composition of the scars
  - o Patients in the active arm had a mean reduction in collagen<sup>1</sup> of 30% compared to placebo after three months treatment. (p<0.01)
- Longer study required to show appearance and physical improvements
  - No significant differences in the overall POSAS<sup>2</sup> score were seen between active and placebo groups after three months of treatment.



# PXS-6302 Phase 1c Trial (Solaria 2); Expert review

- "Exploratory clinical study has significantly enhanced our understanding of the role of LOX enzymes in scarring and the scar process itself."
- "PXS-6302 leads directly to an unprecedented change to the scar composition that we have not seen with any other form of treatment. We estimate that up to 50% of the excess collagen in these patients' scars has been removed."
- "While the length of this Phase 1c safety study was not sufficient to change the appearance of an established scar the remodelling process will be ongoing and I'm confident we would see an improvement in scar appearance and physical characteristics if we observed them for longer."



Professor Fiona Wood
Burns Service of Western Australia
Director of the Burn Injury Research Unit
University of Western Australia

# Phase 1c Established Scar Trial (Solaria 2); Next steps

- Positive data from Solaria 2 trial leads to extension of collaboration with Professor Wood's UWA team
- Wide vista of potential skin fibrosis indications opened up for clinical development. For example:
  - Younger scars
  - Scar prevention post surgery
  - Keloids
  - Dupuytren's
  - Surgical adhesions
- Further update on plans for skin scarring franchise 2H 2023

# **Upcoming News Flow**



### **Upcoming News Flow**

### Five trials to deliver near term value

Pipeline creates multiple opportunities in high value markets

	Indication	Addressable market (US\$)	Trial design	# patients	Status	Data
PXS-5505	Myelofibrosis (MF)	\$1 billion	Phase 2 open label 6 month study in JAK intolerant / ineligible myelofibrosis patients	24	Recruiting	Final interim data released July 2023
			Phase 2 open label 6 month study in JAK intolerant / ineligible myelofibrosis patients	TBD	First Patient 2H 2023	TBD
PXS-6302	Modification of established scars	\$3.5 billion	Phase 1c 3 month placebo controlled study in patients with established scars (>1 year old)	50	Reported	Top line results released May 2023
PXS-	Scar prevention	\$3.5 billion	Phase 1c 3 month placebo controlled study in patients with scarring subsequent to a burns injury	50	First patient 2H 2023	2024
PXS-4728	Isolated REM sleep behaviours disorder (iRDB) and neuro inflammation	\$3.5 billion	Phase 2 double blind, placebo controlled study in patients with iRBD	40	First patient Q3 2023	H1 2025

### **Upcoming News Flow**

### **News flow**

Recent and anticipated news flow

Strong and growing pipeline with advancement in studies expected to provide value inflection points



#### Q1 2023

- Pharmaxis strengthens Board with two new appointments
- PXS-5505 publication by KOL in hematological cancer myelodysplastic syndrome



#### Q2 2023

- PXS-5505: Encouraging FDA feedback on plans to progress to JAK inhibitor combination study
- LOX topical drug PXS-6302 top line data from established scars study
- PXS-5505 myelofibrosis monotherapy study: significant data update



#### H<sub>2</sub> 2023

- PXS-5505 phase 2 myelofibrosis study add on to JAK inhibitor commences recruitment
- Pan-LOX scar treatment and prevention clinical development update and trial initiation
- PXS-4728 iRBD / neuro inflammation study commences recruitment
- PXS-5505 phase 2a myelofibrosis study completed and reports safety and efficacy data at ASH



# **Shareholders & cash**



Financial Information	18 July 23
ASX Code	PXS
Share price	\$0.054
Liquidity (turnover last 12 months)	91m shares
Market Cap	A\$40m
Cash balance (31 March 2023)	A\$15m
Enterprise value	A\$25m
Clinical development program supported by: • R&D tax credits (PXS 2022 claim was \$5m) • Strategy of partnering deals with pipeline asse	ets

Institutional Ownership	30 June 23	
BVF Partners LP	14%	
Karst Peak Capital Limited	12%	
D&A Income Limited	11%	
Platinum Investment Management Limited	8%	
Regal Funds Management Pty Ltd	5%	
Total Institutional Ownership	50%	







# pharmaxis

developing breakthrough treatments for fibrosis and inflammation

Pharmaxis Ltd ABN 75 082 811 630 www.pharmaxis.com.au





Contacts
Gary Phillips
Chief Executive Officer
gary.phillips@pharmaxis.com.au

David McGarvey Chief Financial Officer david.mcgarvey@pharmaxis.com.au