## **Company presentation**

June 2020

NASDAQ: ASLN TPEx: 6497



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### Management team with global development experience

Position	Experience	
Dr Carl Firth CEO	AstraZeneca Head of New Portfolio (China) Head of BD (Asia)	Bank of America Merrill Lynch Head of Asia Healthcare Banking
Stephen Doyle CBO	Boehringer Ingelheim VP Specialty Care & Diabetes (China)	SANOFI 🎝 VP Oncology (China)
Dr Bertil Lindmark Acting CMO	AstraZeneca Head of Development, R&I Head of Development, Japan	Global Head of R&D CSO
Alison Ward VP R&D	AstraZeneca	Sinnovation for patient care Biologics process development
Kiran Asarpota VP Finance	GLOBAL BRANDS GROUP GROUP Finance Director	
Ben Goodger General Counsel	Senior Partner and Head of IP	EDWARDS WILDWAN Partner and Head of IP



## Clinical-stage immunology & oncology biopharma developing innovative treatments to transform the lives of patients

Portfolio led by ASLAN004, a monoclonal antibody targeting IL-13R $\alpha$ 1, that has the potential to be best-in-disease for atopic dermatitis and asthma

Programs	Discovery	Preclinical	Phase 1	Phase 2	Anticipated milestones
Immunology					
ASLAN004	Atopic dermatit	is			<ul> <li>MAD interim data 2H 2020</li> <li>MAD completion 1H 2021</li> </ul>
IL-13Rα1 inhibitor	Asthma				
Oncology					
<b>ASLAN003</b> DHODH inhibitor	AML				
Discovery					
AhR antagonist <sup>1</sup>	Oncology				

1 Aryl hydrocarbon receptor, or AhR, program is being developed in an ASLAN majority-owned joint venture



## ASLAN004: the potential to lead a \$21B market

- Atopic dermatitis (AD) is a large and growing market predicted to exceed \$21 billion by 2027
- Biologics are changing the treatment paradigm in atopic dermatitis
- Dermatologists indicate the need for new, differentiated therapies

<b>x</b>	

- ASLAN004 is a first-in-class inhibitor of the IL13 receptor and has the potential to be a best-in-disease therapy for AD
- Potential for improved efficacy, fewer adverse events and monthly dosing
- ASLAN004 is the only drug besides *dupilumab* to provide dual IL4/IL13 blockade, a validated approach which is becoming standard of care for moderate-to-severe AD
- Potential in asthma and other indications driven by allergic inflammation

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- Phase 1 SAD in healthy volunteers complete. No significant adverse events noted to date. Profile may allow for monthly dosing
- Currently recruiting second cohort in MAD / PoC study. Early efficacy data encouraging. Expected completion in 1H 21
- Phase 2B to start in 2021



## Atopic dermatitis is a chronic disease that can severely impact quality of life

**Total AD prevalent cases** 



- Atopic dermatitis (AD) is a chronic inflammatory skin condition and the most common form of eczema
- Characterised by red inflamed skin and severe daytime and night-time itching
- Over 200 million AD patients worldwide
  - Prevalence estimated at 1-3% of adults worldwide
  - Up to 50% are moderate-tosevere patients



### Market is being driven by biologics and systemic therapies

- There are few safe and effective treatments for moderate-to-severe AD
- Treatment traditionally focused on topical corticosteroids
- Systemic steroids associated with safety risks
- Dupilumab (approved in 2017) is the only biologic therapy available today
- *Dupilumab* has set a new standard for treating AD, but physicians are looking for better safety, improved efficacy, dosing regimen and convenience

	Efficacy	Safety	Dosing	Convenience	Possible use (if approved)
Target profile	+++	+++	+++	+++	
Dupilumab (IL4R)	++	++	+	+	Standard of care for systemic therapy
IL13 eg lebrikizumab	+	+++	+/++	?	When dupi can't be tolerated?
Pan-JAK eg baricitinib	+++	-	++	+++	
JAK-selective eg abrocitinib	+++	-/+	++	+++	When dupi fails?

Comparison of treatment options approved and in development for moderate-to-severe AD



### AD is where the psoriasis market was 15 years ago

- The US psoriasis market grew from \$1bn to \$10bn in 15 years driven by the launch of biologics
- The AD landscape is poised for similar growth driven also by novel biologics
- AD market expected to exceed \$21 billion by 2027



Source: Decision Resources

# *Dupilumab* has advanced the standard of care for AD, but a significant unmet need remains

- Launch of *dupilumab* in 2017 helped drive a large market for systemic AD therapy
- Dupilumab establishing dual blockade of IL4/IL13 biologic therapy as the new standard of care
- Sanofi intends to grow sales to over \$11B
- However, there remains a significant unmet need
  - Only 35% of patients treated with *dupilumab* achieved an optimal response<sup>1</sup>
  - Conjunctivitis common and can lead to treatment discontinuations
  - Opportunity to improve on biweekly dosing regimen



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### Real world studies suggest conjunctivitis and eosinophilia may be more common and lead to treatment discontinuations

- In a review of 29 clinical studies, rates of conjunctivitis varied from 9 to 22%<sup>1</sup>
- Data from real world studies suggest rates in clinical practice may be much higher<sup>2,3</sup>
- Onset of conjunctivitis can be from weeks to months<sup>1</sup>
- In a recent retrospective study of 241 patients on *dupilumab*:
  - 38% of patients experienced conjunctivitis and 56% eosinophilia
  - 17% patients discontinued treatment mostly because of safety

Retrospective study of 241 AD patients taking dupilumab<sup>2</sup>

Adverse event	Patier	nts (%)
At least 1 AE	171	(71)
Noninfectious ophthalmologic	107	(49)
Conjunctivitis	84	(38)
Ocular pruritus	52	(24)
Blepharitis	31	(14)
Xerophthalmia	27	(12)
Keratitis	14	(6)
Eosinophilia	100	(56)
Over 3x normal <sup>4</sup>	28	(30)

Reasons for discontinuation in 42 patients:



- 1 Agnihotri et al, 2019. Drugs in R&D. 19:311
- 2 Faiz et al, 2019. JAAD. 81:143. 241 moderate-to-severe AD pts collected in 2017 and 2018
- 3 Wollenberg et al, 2018. J Allergy Clin Imm Pract. 6:1778
- 4 Over 1500 cells/mm<sup>3</sup>



## ASLAN004 is a first-in-class IL-13R antibody that has the potential to be best-in-disease

#### Target profile:

A drug that can deliver better efficacy over current standard of care

A drug that addresses physician concerns on safety with lower rate of discontinuation



A drug that allows monthly dosing for patients improving convenience and compliance

A drug with greater storage flexibility allowing it to be stored at room temperature



## ASLAN004 is the only drug besides *dupilumab* to provide dual IL4 / IL13 blockade



ASLAN004 blocks the formation of the Type II receptor, preventing signalling through both IL4 and IL13



# Dual-blockade (receptor targeting) appears clinically to be more effective

IL4/IL13 receptor targeting			
ASLAN004 (ASLAN)	IL13Ra1	Phase 1 / POC in atopic dermatitis	
Dupilumab (Sanofi / Regeneron)	IL4Rα	Approved in atopic dermatitis and allergic asthma	
IL4/IL13 ligand targeting			
<i>Lebrikizumab</i> (Eli Lilly / Dermira)	IL13	Discontinued in asthma, phase 3 in atopic dermatitis	
Tralokinumab (Leo)	IL13	Discontinued in asthma, phase 3 in atopic dermatitis	
Altrakincept (Amgen)	IL4	Discontinued	
Pascolizumab (GSK)	IL4	Discontinued	
Receptor targeting		Ligand targeting	
% change in EASI			
80%			
60%	<b></b>	∕	
40%	∆289	°∆17%	
20%			
2070			
0% 300mg 300mg 300mg 300mg	250m	g 250mg 300mg	
q2w q2w q2w	q4w	q2w q2w + TCS	
Dupi SOLO1 Dupi SOLO2 Dupi ph (wk 16) (wk 16) (wk 16	2b	Lebri ph2b Tralo ph2b (wk 16) (wk 12)	

## *Dupilumab*-associated conjunctivitis may be driven by inhibition of Type I receptor, which ASLAN004 does not bind



- *Dupilumab* blocks the Type I receptor
- This may drive  $T_H^2$  to  $T_H^1$  polarisation
- T<sub>H</sub>1 cells product interferon gamma, which can lead to apoptosis of goblet cells
- This could lower the production of mucin and lead to development of dry eye and conjunctivitis

ASLAN004 binds more strongly to receptor than dupilumab relative to its respective ligand

Receptor	Ligand	K <sub>D</sub> (nM)	Comments
IL-13Rα1	IL-13	30 <sup>1</sup>	ASLAN004 has a 60 fold higher
IL-13Rα1	ASLAN004	0.5	affinity for receptor than IL-13
IL-4Rα	IL-4	0.1 <sup>1</sup>	Dupilumab only has a 3 fold higher
IL-4Rα	Dupilumab	0.03	affinity for receptor than IL-4







ASLAN004 offers a greater affinity difference between ligand and receptor binding which may translate to lower required concentration in vivo and may provide improved dosing frequency and efficacy



## Phase 1 SAD study in healthy volunteers completed

- Well tolerated at all doses when administered IV and subcutaneous (SC)
  - No adverse events that led to discontinuations, no significant injection site reactions
  - SC now used in on-going studies
- Analysis of downstream mediators including pSTAT6 demonstrated complete inhibition within one hour of dosing with IV and maintained for more than 29 days
- PK profile may allow for monthly dosing
- Concentration required to completely inhibit signal transduction via the receptor was over an order of magnitude lower than that of existing therapies



## ASLAN004 well-tolerated at all dose levels

Drug-related adverse event	N = 44				
	Any	grade		Severity	
	Ν	(%)	Mild	Moderate	Severe
Decreased appetite	2	5	1	1	0
Alanine aminotransferase increased	1	2	1	0	0
Diarrhoea	1	2	1	0	0
Pyrexia	1	2	1	0	0
Blood lactate dehydrogenase increase	1	2	1	0	0
Weight decrease	1	2	1	0	0
Lymphocyte count decrease	1	2	1	0	0
Headache	1	2	0	1	0
C-reactive protein increase	1	2	1	0	0
Injection site pruritus (mild)	1	2	1	0	0

## ASLAN004 MAD is designed to deliver PoC

- MAD / PoC study testing ASLAN004 in moderate-severe AD patients
- Double-blind, placebo-controlled study
- Patients dosed for 8 weeks with a 12-week recovery period
- Patients recruited from Singapore, opening sites in Australia and US
- Interim unblinded data from cohorts 1-3 expected 2H 20, with completion in 1H 21

Moderate-to- severe atopic dermatitis patients (N = 42)	Cohort 1 – 200mg QW (ASLAN004 N = 6, placebo N = 2)	Expansion cohort –
	Cohort 2 – 400mg QW (ASLAN004 N = 6, placebo N = 2)	Dose 1, 2 or 3 QW (ASLAN004 N = 12,
	Cohort 3 – 600mg QW (ASLAN004 N = 6, placebo N = 2)	placebo N = 6)

Primary endpoints are safety and tolerability

Secondary endpoints include percentage change in EASI score (EASI50, EASI75) pruritus score and IGA, TARC, IgE

Study has 80% power to detect a 39% improvement in the percentage change in EASI score from baseline based on a one-sided 5% significance level



### Early signs of efficacy in low dose cohort

- First patient enrolled on 22 October 2019
- As of 29 November 2019, 6 patients treated in low dose (200mg) cohort
- Currently recruiting 2<sup>nd</sup> dose cohort
- ASLAN004 well-tolerated, with no serious AEs or treatment discontinuation
- 3 patients completed at least 1 month of dosing with average reduction in EASI of 71%
- Maximal efficacy expected at 6 to 8 weeks





## ASLAN004 has the potential to be best-in-disease

IL-13Rα1 inhibitor	ASLAN004 is the only monoclonal antibody targeting IL-13R $\alpha$ 1, and has the potential to be best-in-disease for atopic dermatitis and asthma
Validated pathway	Targets the same pathway and receptor complex (Type II) as <i>dupilumab</i>
Targeting differentiated profile	Potential for improved efficacy, fewer adverse events, monthly dosing
SAD completed	Phase 1 SAD in healthy volunteers completed. No significant adverse events noted to date. Profile may allow for monthly dosing.
MAD/PoC ongoing	Currently recruiting second cohort in MAD / PoC study. Early efficacy data encouraging. Expecting interim data 2H 20, completion in 1H 21
Phase 2b program	Planning to initiate phase 2b program in 2021
Potential in other indications	Potential to pursue development in other indications where <i>dupilumab</i> has proven to be effective







## **Financials**

Shares outstanding Ordinary shares ADS equivalent (5:1)	Overall 190M 38M	NASDAQ: ASLN 59M 12M	TPEx: 6497 (Taiwan) 130M 26M
Net loss	US\$ 3.0M (1Q 20)		
Cash balance	US\$ 16.9M (1Q 20)		
Recent financing	US\$ 14.7M raised in Dec 2019 (NASDAQ follow-on)		

