Company presentation

April 2021

NASDAQ: ASLN



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Management team with global development experience





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

Portfolio led by ASLAN004, a monoclonal antibody targeting IL-13R α 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 IL-13Rα1 inhibitor	Atopic dermatit	is			MAD topline data 3Q21Initiate Phase 2b in 2H21
	Asthma				
ASLAN003 DHODH inhibitor	Autoimmune di	sease			
Discovery					
AhR antagonist ¹	Oncology				

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

ASLAN004: potential first-in-class antibody IL-13R

- We believe dermatologists and patients looking for additional AD treatment options – opportunity to improve on efficacy, safety and dose regimen
- ASLAN004 is a novel, first-in-class antibody targeting IL-13R, blocking both IL-4 and IL-13 signalling through the Type 2 receptor
- Emerging clinical data demonstrate competitive profile with the potential to differentiate over existing therapies
- Robust dose dependent efficacy profile showing improvement compared to placebo across all efficacy endpoints
- Well-tolerated at all doses tested
- Expansion cohort continues to recruit with full study readout expected in 3Q 2021, Phase 2b anticipated to initiate in 2H 2021

Interim data demonstrates a robust and differentiated safety and efficacy profile.



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





- 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



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

- There are few safe and effective treatments for moderate-to-severe AD
- Treatment traditionally focused on topical corticosteroids. Systemic steroids associated with safety risks
- 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 expects to grow sales to over \$11B
- However, there remains a significant unmet need
 - Only 35% of patients treated with *dupilumab* achieved an optimal response¹
 - Conjunctivitis common and can lead to treatment discontinuations
 - Opportunity to improve on biweekly dosing regimen





1 Spherix (2018) Atopic dermatitis ATU study

ASLAN004 is a potential 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 a potential first-in-class IL-13R antibody that selectively blocks the Type 2 receptor



ASLAN004 is the only monoclonal antibody in the clinic targeting IL-13R α 1

- IL-4 and IL-13 are central to triggering allergy and symptoms of atopic dermatitis
- ASLAN004 blocks the Type 2 receptor, preventing signalling through both IL-4 and IL-13

Potential for improved efficacy, safety and dose regimen:

- Selectively targets the Type 2 receptor. Blocking the Type 1 receptor may lead to unwanted effects
- Stronger binding to receptor than *dupilumab* relative to its respective ligand, so pathway inhibition may be maintained at low drug concentrations



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_H2 to T_H1 polarisation
- T_H1 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 _D (nM)	Comments
IL-13Rα1	IL-13	30 ¹	ASLAN004 has a 60 fold higher
IL-13Rα1	ASLAN004	0.5	affinity for receptor than IL-13
IL-4Rα	IL-4	0.1 ¹	Dupilumab only has a 3 fold higher
IL-4Rα	Dupilumab	0.03	affinity for receptor than IL-4



Drug concentration



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



Trial design of MAD / PoC study in moderate-severe AD

- Double-blind, randomised, placebo-controlled study
- Patients dosed weekly via subcutaneous injection for 8 weeks with a 12-week recovery period
- Interim data from cohorts 1 to 3

Moderate-to- severe atopic dermatitis patients (N ≈ 50)	Cohort 1 – 200mg QW (ASLAN004 N ≈ 6, placebo N ≈ 2)	Expansion cohort		
	Cohort 2 – 400mg QW (ASLAN004 N ≈ 6, placebo N ≈ 2)	$\begin{array}{l} 600 \text{mg QW} \\ \text{(ASLAN004 N} \geq 16, \end{array}$		
	Cohort 3 – 600mg QW (ASLAN004 N ≈ 6, placebo N ≈ 2)	placebo N \ge 8)		

Primary endpoints are safety and tolerability

Secondary endpoints include percentage change from baseline in EASI score, pruritus score and IGA, and biomarkers TARC and IgE

Key inclusion criteria:

- Chronic AD present for ≥3 years before screening visit
- EASI score ≥16 at screening and baseline
- IGA score ≥3 (scale of 0 to 4) at screening and baseline
- ≥10% body surface area (BSA) of AD involvement at screening and baseline

Study has 80% power to detect 39% improvement in EASI from baseline, compared to placebo, based on a one-sided 5% significance level



Study recruitment





Selected baseline patient characteristics

Per protocol efficacy evaluable (N=18)	200mg (N=4)	400mg (N=6)	600mg (N=3)	Placebo (N=5)
Age (years)	32.5	28.3	42.0	33.8
Mean EASI score	32.9	30.9	32.5	33.9
Mean BMI	25.8	25.4	24.2	25.4
Patients with IGA 3 / IGA 4	50 / 50 %	83 / 17 %	33 / 67 %	40 / 60 %
Mean BSA	55.5%	59.8%	56.3%	59.8%
Mean peak pruritus NRS score	7.4	7.3	6.4*	7.4

Abbreviations

EASI: Eczema Area and Severity Index

IGA: Investigator Global Assessment

BSA: Body Surface Area

NRS: Numerical Rating Scale

* N=2 as one subject did not have a baseline value



Mean reduction in EASI from baseline (Week 8)





EASI-50 (Week 8)





EASI-75 (Week 8)





EASI-90 (Week 8)





EASI score over time







Patients achieving IGA 0/1 (Week 8)

40%





Mean reduction in peak P-NRS from baseline (Week 8)





* One subject did not have a baseline value

Peak P-NRS over time





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ASLAN004 well-tolerated at all dose levels

Treatment Emergent Adverse Event (TEAE) by category	200mg (N=5)	400mg (N=8)	600mg (N=5)	All doses (N=18)	Placebo (N=7)
Any	5 (100%)	8 (100%)	3 (60.0%)	16 (88.9%)	5 (71.4%)
Related	5 (100%)	6 (75.0%)	2 (40.0%)	13 (72.2%)	5 (71.4%)
Moderate/severe	2 (40.0%)	2 (25.0%)	1 (20.0%)	5 (27.8%)	3 (42.9%)
Serious adverse event (SAE)	0 (0%)	1 (12.5%)	0 (0%)	1 (5.6%)	0 (0%)
Drug-related AEs of interest*:					
Injection site reaction	1 (20.0%)	3 (37.5%)	0 (0%)	4 (22.2%)	2 (28.6%)
Conjunctivitis	0 (0%)	1 (12.5%)	1 (20.0%)	2 (11.1%)	0 (0%)

- There were no drug-related TEAEs that led to discontinuation
- SAE was mild abdominal pain, classified as unlikely related
- Conjunctivitis was classified as mild. Patients had prior history of allergy or allergic conjunctivitis.



Opportunity for better treatment options for patients

	Categories	Dupilumab Ph3 ¹ (300mg QW)		Dupilumab Ph3 ¹ (300mg Q2W)		Lebrikizumab Ph2b ⁵) ⁵
			SOLO2	SOLO1	SOLO2	(250Hg Q2W)		
	Age (years) – mean	39.3 vs 39.5 ²	37.1 vs 37.4 ⁴	39.8 vs 39.5 ²	36.9 vs 37.4 ⁴		38.9 vs 42.2	
	EASI score – mean	33.2 vs 34.5 ²	31.9 vs 33.6 ⁴	33.0 vs 34.5 ²	31.8 vs 33.6 ⁴		25.5 vs 28.9	
Baseline characteristics	Patients with IGA 4	48% vs 49%	47% vs 49%	48% vs 49%	49% vs 49%		29% vs 39%	
	BSA – mean	56% vs 58% ²	52% vs 54% ⁴	55% vs 58% ²	53% vs 54% ⁴		40% vs 47%	
	Pruritis NRS – mean	7.2 vs 7.4 ²	7.5 vs 7.5 ⁴	7.2 vs 7.4 ²	7.6 vs 7.5 ⁴		7.6 vs 7.4	
	Efficacy at	16 weeks	16 weeks	16 weeks	16 weeks	4 weeks	8 weeks	16 weeks
	% change in EASI	-72% vs -38%	-69% vs -31%	-72% vs -38%	-67% vs -31%	-50% vs -25% ⁶	-64% vs -31% ⁶ ·	-73% vs -41% ⁶
	EASI-50	61% vs 25%	61% vs 22%	69% vs 25%	65% vs 22%	NA	NA	81% vs 46%
Efficacy	EASI-75	52% vs 15%	48% vs 12%	51% vs 15%	44% vs 12%	30% vs 3% ⁶	46% vs 17% ⁶	61% vs 24% ⁶
	EASI-90	33% vs 8%	31% vs 7%	36% vs 8%	30% vs 7%	14% vs 1% ⁶	30% vs 4% ⁶	44% vs 11% ⁶
	Patients achieving IGA 0/1	37% vs 10%	36% vs 8%	38% vs 10%	36% vs 8%	14% vs 0% ⁶	31% vs 5% ⁶	45% vs 15% ⁶
	% change in Pruritis NRS	-49% vs -26%	-48% vs -15%	-51% vs -26%	-44% vs -15%	-39% vs -25% ⁶ ·	-46% vs -22% ⁶	-62% vs 7% ⁶
Safety and tolerability	Serious AE	1% vs 5%	3% vs 6%	3% vs 5%	2% vs 6%		3% vs 4%	
	Conjunctivitis	8% vs 2% ³	7% vs 2% ³	12% vs 2% ³	7% vs 2% ³		3% vs 0% ⁷	

Numbers in table refer to drug vs placebo

¹ Simpson et. al, NEJM 1 October 2016 (unless otherwise stated)

² https://clinicaltrials.gov/ct2/show/results/NCT02277743

³ Includes allergic conjunctivitis, conjunctivitis bacterial and conjunctivitis viral as reported in the supplementary appendix of the source document

- ⁴ https://clinicaltrials.gov/ct2/show/results/NCT02277769
- ⁵ Guttman-Yassky et al, JAMA Dermatology, 26 Feb 2020 (unless otherwise stated)
- ⁶ Lebrikizumab Program Update, 17 October 2019 by Dermira

⁷ Includes conjunctivitis, conjunctivitis bacterial and conjunctivitis allergic as reported in the source document

Summary of positive interim data and next steps

Interim analysis summary

- 74% average reduction in EASI from baseline at therapeutic doses (400mg and 600mg) after 8 weeks
 - 89% of patients achieved EASI-50
 - 56% achieved EASI-90
- Data supportive of ASLAN004's potential as a novel, first-in-class antibody targeting IL-13R with differentiated efficacy and safety profile

Next steps

- Expansion cohort ongoing
 - Targeting the recruitment of at least 24 patients
 - Readout expected 3Q 2021
- Phase 2b planning underway, expected to initiate in 2H 2021
- Prioritising additional indications for potential new studies in 2H 2021







ASLAN003 is an orally active, potent inhibitor of DHODH



- Rate-limiting step in the *de novo* synthesis of pyrimidines by the mitochondria
- Inhibition of DHODH reduces the pyrimidine pool used by cells with high metabolic activity
- Normal cells utilising the pyrimidine salvage pathway remain unaffected
- Reduces pro-inflammatory cytokines
- Inhibits proliferation of metabolically active cells such as:



Immune cells which may attack native tissue (autoimmune \rightarrow eg MS, RA)



Cells affected by infectious agents (infectious disease \rightarrow eg COVID-19)



Dysfunctional or malignant cells (**oncology** \rightarrow eg AML/MDS)



ASLAN003 has the potential to be best-in-class for autoimmune disease

- DHODH an effective target in MS and RA treatment (Aubagio, Arava)
 - Aubagio \$2.2B global sales in 2019, part of a \$23B global MS market 1
- ASLAN003 was designed to be more potent and to address the toxicities associated with first generation inhibitors (*leflunomide, teriflunomide*)
 - Superior *in vitro* potency as compared with other DHODH inhibitors
 - Selective against a panel of 195 enzymes, ion channels and receptor binding assays
 - In vitro studies demonstrated ASLAN003 has lowest potential for hepatotoxicity out of 6 approved and clinical stage DHODH inhibitors
- Active in the multiple sclerosis EAE model and rheumatoid arthritis AIA model
- Well tolerated in 119 subjects in Phase 1 and Phase 2 clinical trials
- PK profile suitable for once-daily dosing

Assay used to measure IC ₅₀	ASLAN003 (µM)	<i>Teriflunomide</i> (μM)
Enzymatic DHODH inhibition	0.035	1.1
Human PBMC proliferation inhibition	1.4	46
IFN_{Y} inhibition in human whole blood	2.5	259







Financials

Ticker	NASDAQ: ASLN		
Shares outstanding ¹	69.2M	(as of December 31, 2020 pro forma)	
Net operating cash used	US\$ 5.1M	(Q4 2020)	
Cash balance	US\$ 112.2M	(as of December 31, 2020 pro forma)	
Recent financings	US\$ 69.0M raised in Mar 2021 via public offering US\$ 18.0M raised in Feb 2021 via private placement		

