

# Company presentation

December 2019

NASDAQ: ASLN  
TPEX: 6497



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All statements other than statements of historical fact are forward-looking statements. The words “believe,” “may,” “might,” “could,” “will,” “aim,” “estimate,” “continue,” “anticipate,” “intend,” “expect,” “plan,” or the negative of those terms, and similar expressions that convey uncertainty of future events or outcomes are intended to identify estimates, projections and other forward-looking statements. Estimates, projections and other forward-looking statements speak only as of the date they were made, and, except to the extent required by law, the Company undertakes no obligation to update or review any estimate, projection or forward-looking statement.



# Clinical-stage biopharma with oncology and immunology focus

Platform leverages Asia clinical centres combined with US/EU centres to accelerate clinical development

Programs	Discovery	Preclinical	Phase 1	Phase 2	Anticipated milestones
<b>ASLAN004</b> <i>IL-4/IL-13</i> <i>Receptor inhibitor</i>	Atopic dermatitis				<ul style="list-style-type: none"> <li>• MAD interim data early 2020</li> <li>• MAD completion 2H 20</li> </ul>
	Asthma				
<b>ASLAN003</b> <i>DHODH inhibitor</i>	AML				
<b>Varlitinib</b> <b>(ASLAN001)</b> <i>Pan-HER inhibitor</i>	Gastric cancer (2 <sup>nd</sup> line)				
	Neo-adj breast cancer				
	Hepatocellular carcinoma (2 <sup>nd</sup> line)				
	Biliary tract cancer (1 <sup>st</sup> line)				

## Discovery programs

<b>AhR antagonist<sup>1</sup></b>	Oncology				
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<sup>1</sup> Aryl hydrocarbon receptor, or AhR, program is being developed in an ASLAN majority-owned joint venture with Bukwang Pharmaceutical Co., Ltd.  
 Investigator initiated trial



# Asia offers a unique opportunity to accelerate global clinical development

Where the diseases are more prevalent

Cancer prevalence	US	Asia-Pac	Difference in prevalence
Biliary tract	12,601	200,968	2.8 x
Gastric	32,076	1,027,691	5.6 x
Liver	27,479	422,635	2.7 x
Nasopharyngeal	6,072	112,790	3.2 x

Where there are fewer competing clinical trials

Trials per M capita	US	Asia-Pac (ex-JP)	Difference in density
All diseases	367	18	20 x
AML	5.35	0.03	167 x
Atopic dermatitis	1.03	0.02	62 x

- 1 Gastric, liver and nasopharyngeal cancer: as of 2012, based on Globocan (2012); Bray et al (2013), Estimates of global cancer prevalence for 27 sites in the adult population in 2008.
- 2 Biliary tract cancer: as of 2016, based on Edison Investment Research, "Novel treatments for worldwide unmet needs," dated November 8, 2017, surveying Randi et al (2008), Epidemiology of biliary tract cancers: an update; Bridgewater et al (2014), Guidelines for the diagnosis and management of intrahepatic cholangiocarcinoma.
- 3 In this table, Asia-Pac refers to only China, Japan, Philippines, Singapore, South Korea, Thailand and Vietnam.
- 4 Clinical trial density is defined as number of trials per one million population.



# Management team with global development experience

Position		Experience	
<b>Dr Carl Firth</b> CEO		 Head of New Portfolio (China) Head of BD (Asia)	 Head of Asia Healthcare Banking
<b>Dr Mark McHale</b> CDO Head of R&D		 Head of Molecular Sciences, R&I Head of Early Asthma Portfolio	
<b>Dr Bertil Lindmark</b> Acting CMO		 Head of Development, R&I Head of Development, Japan	 Global Head of R&D CSO
<b>Stephen Doyle</b> CBO		 VP Specialty Care & Diabetes (China)	 VP Oncology (China)
<b>Kiran Asarpota</b> VP Finance		 Group Finance Director	



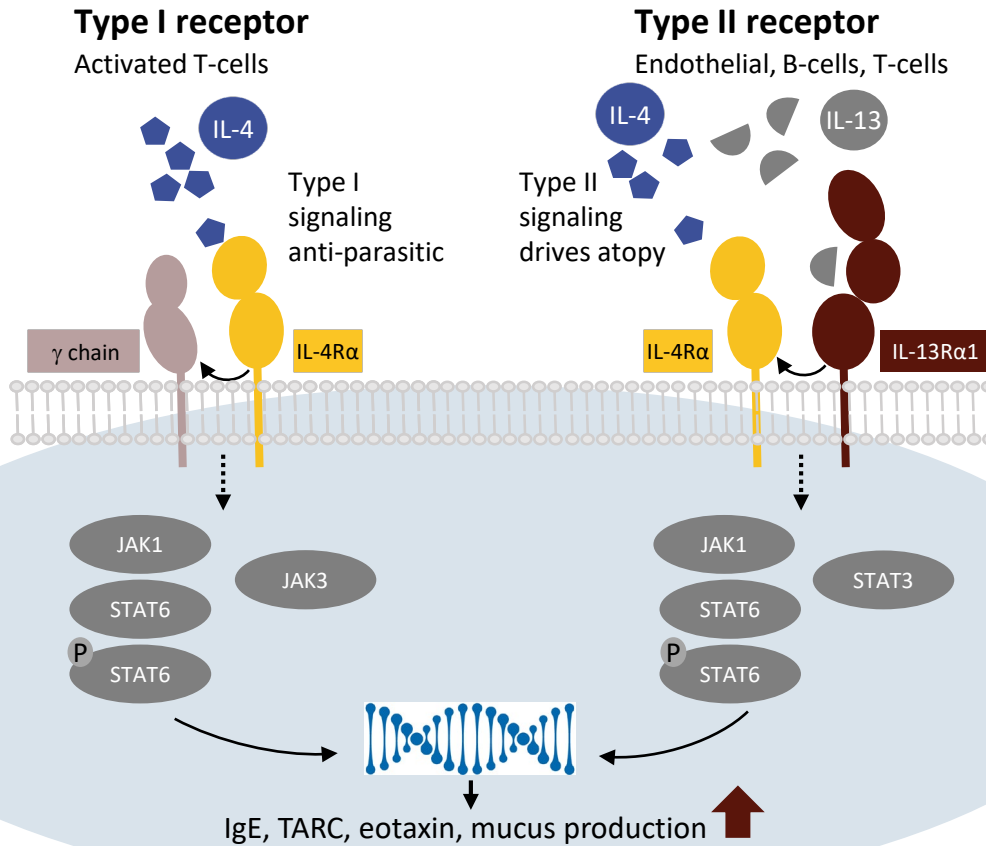
ASLAN004



# ASLAN004 blocks signaling through IL-4 and IL-13

- ASLAN004 targets the IL-13 receptor  $\alpha 1$  subunit
- Blocks same pathways responsible for allergic inflammation as *dupilumab*

**Dupilumab**  
Binds IL-4R $\alpha$ ,  
blocking the  
Type I and Type II  
receptors



**ASLAN004**  
Binds IL-13R $\alpha 1$ , blocking  
the Type II receptor,  
which is responsible for  
allergic inflammation



# Receptor targeting is more effective than ligand targeting

## IL4/IL13 receptor targeting

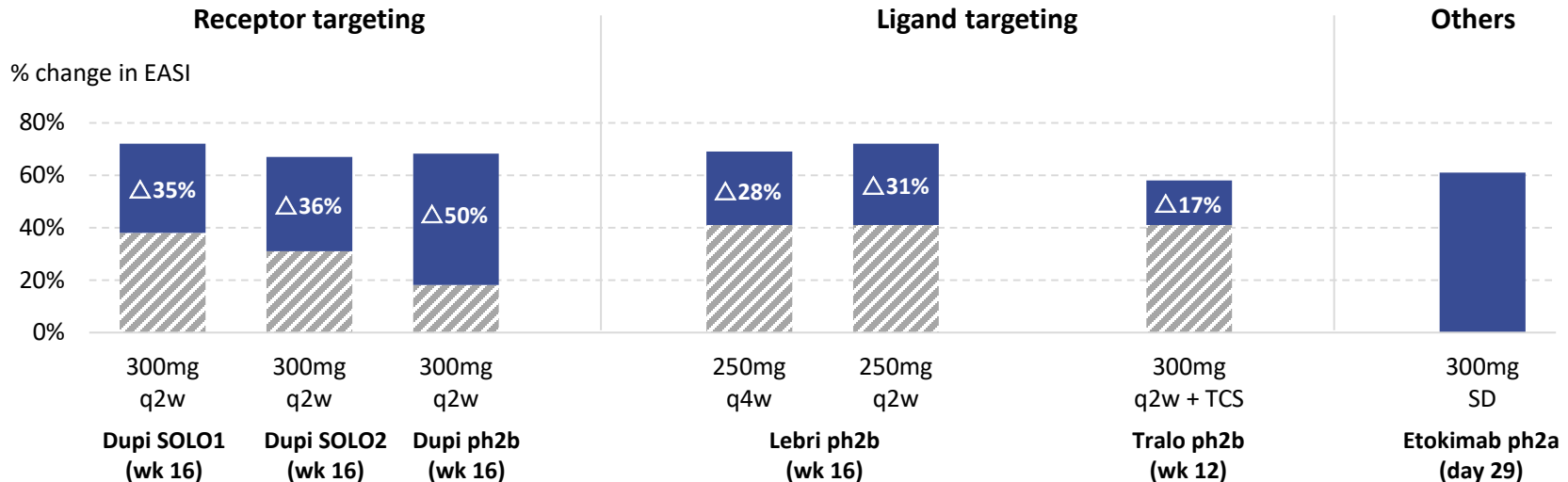
<b>ASLAN004</b>	<b>IL-13R<math>\alpha</math>1</b>	<b>Phase 1 / POC in atopic dermatitis</b>
<i>Dupilumab</i>	IL-4R $\alpha$	Approved in atopic dermatitis and allergic asthma

## IL4/IL13 ligand targeting

<i>Lebrikizumab</i>	IL-13	Discontinued in asthma, phase 3 in atopic dermatitis
<i>Tralokinumab</i>	IL-13	Discontinued in asthma, phase 3 in atopic dermatitis
<i>Altrakincept</i>	IL-4	Discontinued
<i>Pascolizumab</i>	IL-4	Discontinued

## Other targets

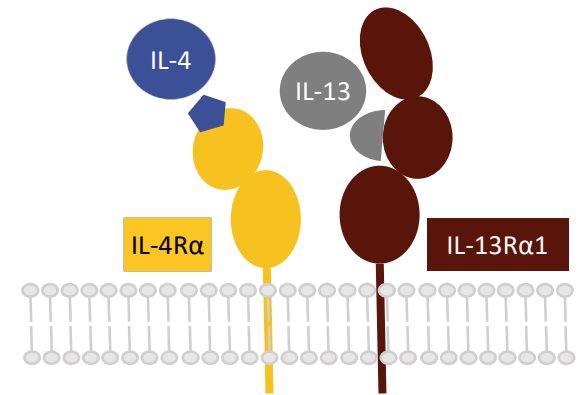
<i>Etokimab</i>	IL-33	Discontinued in atopic dermatitis
MOR106	IL-17C	Discontinued in atopic dermatitis





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

Receptor	Ligand	Kd (nM)	Comments
IL-13R $\alpha$ 1	IL-13	30	ASLAN004 has a 60 fold higher affinity for receptor than IL-13
IL-13R $\alpha$ 1	ASLAN004	0.5	
IL-4R $\alpha$	IL-4	0.1	<i>Dupilumab</i> only has a 3 fold higher affinity for receptor than IL-4
IL-4R $\alpha$	<i>Dupilumab</i>	0.03	



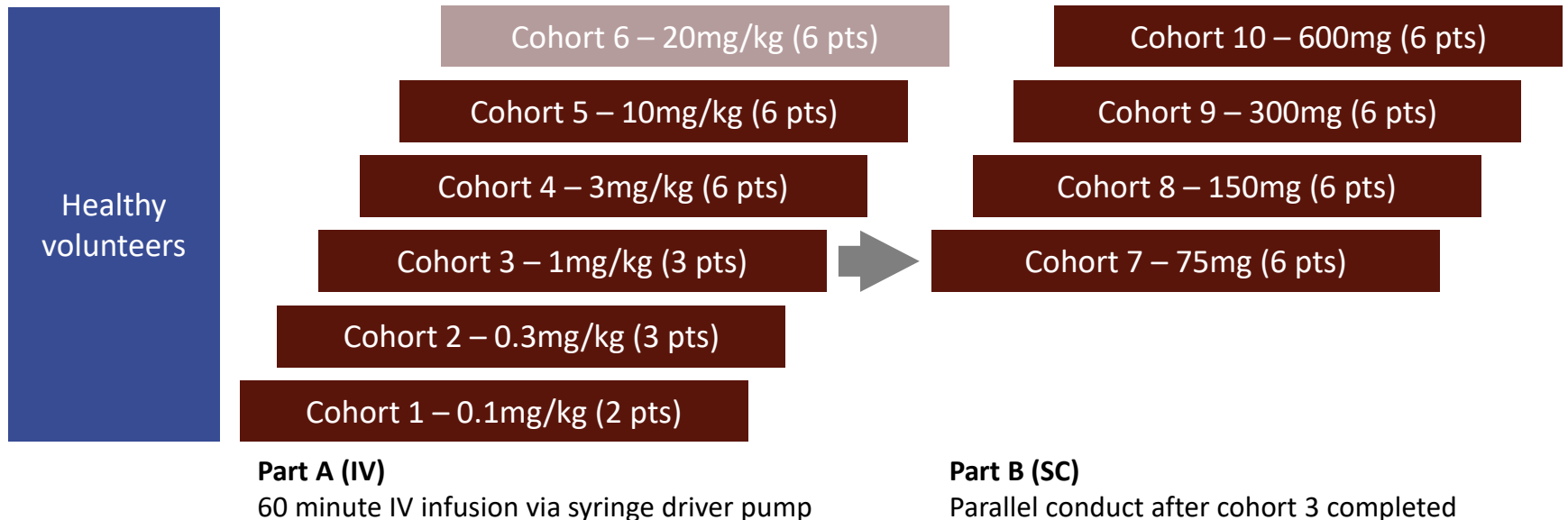
ASLAN004 offers a greater affinity difference between ligand and receptor binding which may translate to lower required concentration *in vivo*



# Phase 1 SAD study in healthy volunteers completed

- Well tolerated at all doses when administered IV and subcut
  - No adverse events that led to discontinuations, no significant injection site reactions
- 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
- Trough level required to completely inhibit signal transduction via the receptor was over an order of magnitude lower than that of existing therapies

Cohort 6 not actioned as long PD effect >29 days achieved at 10 mg/kg



# ASLAN004 well-tolerated at all dose levels

Drug-related adverse event	N = 44				
	Any grade		Severity		
	N	(%)	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 atopic dermatitis patients
- Expected to complete in 2H 20
- Double-blind, placebo controlled study
- Patients dosed for 8 weeks with a 12 week recovery period

Moderate-to-severe atopic dermatitis patients (N = 42)

Cohort 1 – Dose 1 QW  
(ASLAN004 N = 6, placebo N = 2)

Cohort 2 – Dose 2 QW  
(ASLAN004 N = 6, placebo N = 2)

Cohort 3 – Dose 3 QW  
(ASLAN004 N = 6, placebo N = 2)

Expansion cohort –  
Dose 1, 2 or 3 QW  
(ASLAN004 N = 12,  
placebo N = 6)

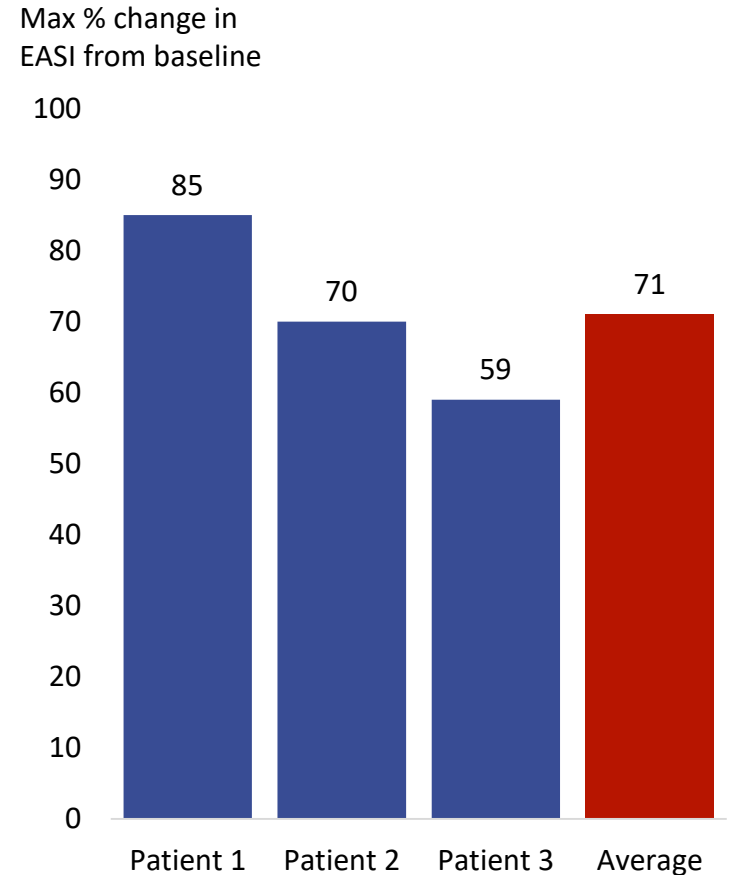
Primary endpoints are safety and tolerability

Secondary endpoints include percentage change in EASI score, EASI50, EASI75, pruritus score and IGA



# 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
- 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
- Data monitoring committee will meet in late December, after which 2<sup>nd</sup> dose cohort expected to open
- Additional interim data from the first 2 cohorts expected in early 2020



# ASLAN004 is a first-in-class IL-13R antibody and has the potential to be superior to *dupilumab*

	<i>Dupilumab</i>	ASLAN004
Efficacy	<ul style="list-style-type: none"><li>• Blocks signaling through IL-4 and IL-13</li><li>• High steady state concentration needed for full target inhibition</li></ul>	<ul style="list-style-type: none"><li>• Blocks signaling through IL-4 and IL-13</li><li>• Only 1mg/l needed for full target inhibition</li></ul>
Dosing	<ul style="list-style-type: none"><li>• Dosed 300mg every 2 weeks</li></ul>	<ul style="list-style-type: none"><li>• Potential for 4 weekly dosing</li><li>• Complete inhibition of pSTAT6 to 29 days after a single IV dose</li></ul>
Safety	<ul style="list-style-type: none"><li>• Conjunctivitis reported between 25% and 50% in clinical practice</li><li>• Injection site reactions common potentially due to formulation</li></ul>	<ul style="list-style-type: none"><li>• No conjunctivitis seen to date</li><li>• No significant injection site reactions seen to date</li></ul>
Stability	<ul style="list-style-type: none"><li>• Cannot be stored above 25°C</li></ul>	<ul style="list-style-type: none"><li>• Over 9 months stability at 25°C</li><li>• Greater flexibility for storage and travel</li></ul>

1 Reported 25-50% conjunctivitis: Wollenberg et al (2018), Conjunctivitis occurring in atopic dermatitis patients treated with dupilumab-clinical characteristics and treatment.

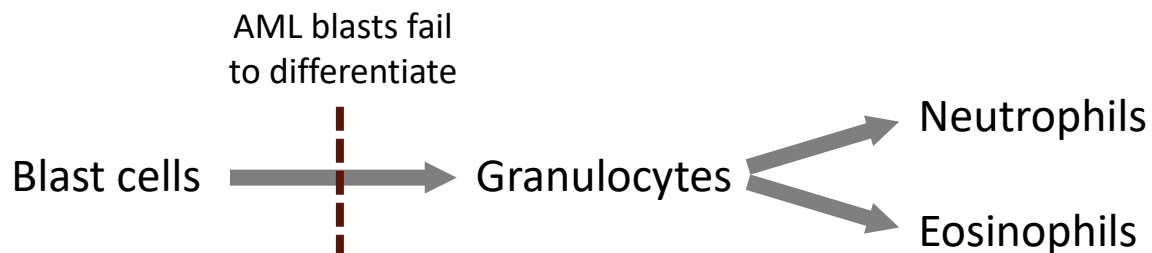


ASLAN003



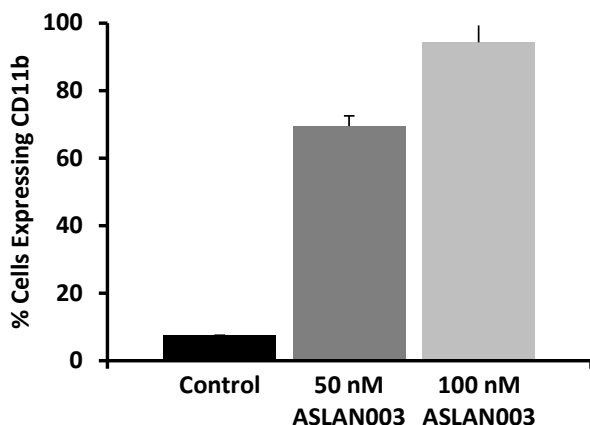
# ASLAN003 is an orally active, potent inhibitor of DHODH

DHODH controls the rate-limiting step in the synthesis of pyrimidines and contributes to the production of ATP

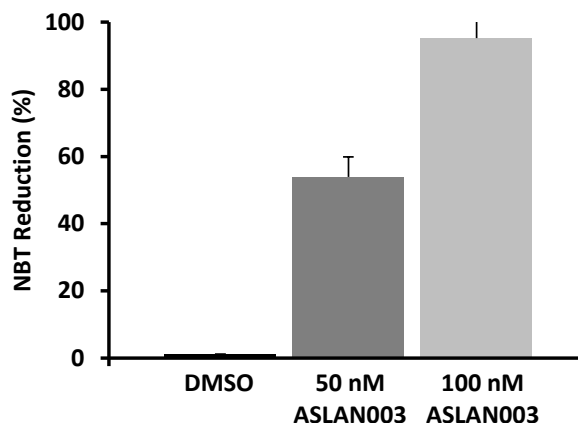


ATRA differentiates blasts in up to 15% of AML patients with >90% CR

Upregulation of CD11b in AML blast cell line THP-1 with ASLAN003



Formation of active granules in AML blast cell line THP-1 with ASLAN003



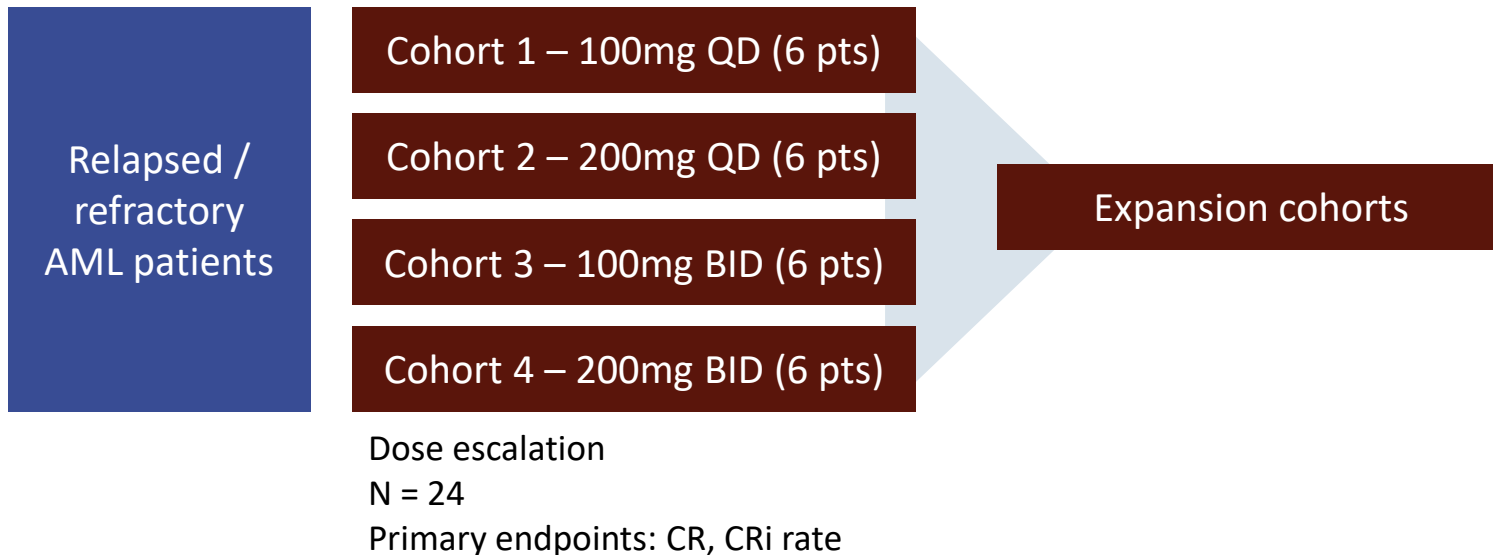
ASLAN003 promotes differentiation in multiple cell lines that are unresponsive to ATRA. This differentiation is mediated via DHODH.





# Phase 2 in relapsed / refractory AML

- 24 patients enrolled in dose escalation cohorts
- 10 patients on treatment for more than 2 months (efficacy evaluable), of which:
  - Fall in peripheral blood blast cells in all evaluable patients (median >50%)
  - 1 PR (reduction in bone marrow blast cells from 54% at baseline to 24%)
  - Evidence of differentiation syndrome seen in some patients
- Well-tolerated, most commonly occurring adverse events: leukocytosis (2 patients grade 3-4), nausea, abd pain and rash



# Other portfolio



# Varlitinib is a potent oral, reversible pan-HER inhibitor

Programs	Discovery	Preclinical	Phase 1	Phase 2
<b>Varlitinib</b> <b>(ASLAN001)</b> <i>Pan-HER inhibitor</i>	Gastric cancer (2 <sup>nd</sup> line)			
	Neo-adj breast cancer			
	Hepatocellular carcinoma (2 <sup>nd</sup> line)			
	Biliary tract cancer (1 <sup>st</sup> line)			

## Competitive efficacy

Increase activity over standard of care: 44% ORR in 1<sup>st</sup> line BTC, 60% ORR in 2<sup>nd</sup> line HER2+ and 60% pCR in neoadj breast cancer.

## Differentiated safety

Substantially lower GI tox compared to other pan-HER inhibitors. 4% drug-related grade 3/4 diarrhoea across all studies.

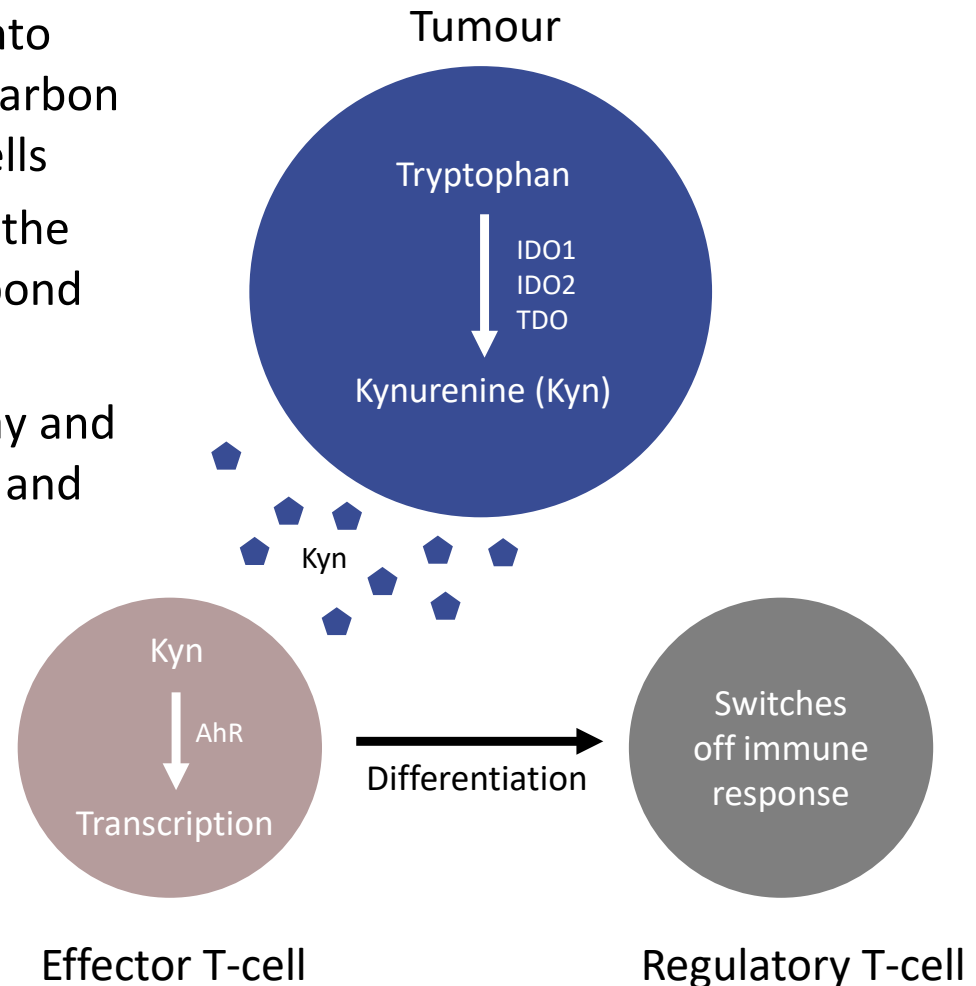
## Efficacy in subgroups of BTC

Pre-specified subgroups in phase 2 TreeTopp study and China Jadetree study show possible benefit. Further analysis ongoing



# We are developing AhR antagonists in our majority-owned subsidiary JAGUAHR Therapeutics

- Tumours break down tryptophan into Kyn, which binds to the aryl hydrocarbon receptor (AhR) and suppresses T-cells
- The Kyn pathway is used to ensure the immune system does not over-respond to threats
- Tumours have hijacked this pathway and frequently overexpress IDO1, IDO2 and TDO
- Established a JV with Bukwang, who are investing up to \$5M to deliver IND-ready compounds
- ASLAN retains buy-back option on assets



# Financials



# Financials

<b>Exchange / ticker</b>	<b>US – NASDAQ: ASLN Taiwan – TPEX: 6497</b>
Shares outstanding	190.0M (equivalent to 38.0 ADSs)
Net loss	US\$ 5.2M (for 3Q 19)
Cash balance	US\$ 10.4M (unaudited, end of Nov 19)
Recent financing	US\$ 14.7M raised in Dec 2019 (NASDAQ follow-on)

