UNITED STATES
SECURITIES AND EXCHANGE COMMISSION
Washington, DC 20549

FORM 8-K

CURRENT REPORT
Pursuant to Section 13 or 15(d) of
The Securities Exchange Act of 1934

Date of Report (Date of earliest event reported): December 2, 2019

Aclaris Therapeutics, Inc.
(Exact name of registrant as specified in its charter)

Delaware	**001-37581**	**46-0571712**
(State or other jurisdiction of incorporation)	(Commission File Number)	(IRS Employer Identification No.)

640 Lee Road, Suite 200
Wayne, PA 19087
(Address of principal executive offices, including zip code)

(484) 324-7933
(Registrant's telephone number, including area code)

N/A
(Former name or former address, if changed since last report)

Check the appropriate box below if the Form 8-K filing is intended to simultaneously satisfy the filing obligation of the registrant under any of the following provisions:

[ ] Written communications pursuant to Rule 425 under the Securities Act (17 CFR 230.425)

[ ] Soliciting material pursuant to Rule 14a-12 under the Exchange Act (17 CFR 240.14a-12)

[ ] Pre-commencement communications pursuant to Rule 14d-2(b) under the Exchange Act (17 CFR 240.14d-2(b))

[ ] Pre-commencement communications pursuant to Rule 13e-4(c) under the Exchange Act (17 CFR 240.13e-4(c))

Securities registered pursuant to Section 12(b) of the Act:

Title of Each Class:	Trading Symbol(s)	Name of Each Exchange on which Registered
Common Stock, $0.00001 par value	ACRS	The Nasdaq Stock Market, LLC

Indicate by check mark whether the registrant is an emerging growth Company as defined in Rule 405 of the Securities Act of 1933 (§230.405 of this chapter) or Rule 12b-2 of the Securities Exchange Act of 1934 (§240.12b-2 of this chapter).

Emerging growth company ☑

If an emerging growth company, indicate by check mark if the registrant has elected not to use the extended transition period for complying with any new or revised financial accounting standards provided pursuant to Section 13(a) of the Exchange Act. ☑

Item 7.01 Regulation FD Disclosure.

On December 4, 2019, management of Aclaris Therapeutics, Inc. (the "***Company***") will present a company overview at the Evercore ISI 2nd Annual HealthCONx Conference in Boston, Massachusetts. The presentation will include a slide presentation. A copy of this slide presentation is furnished as Exhibit 99.1 to this Current Report on Form 8-K.

In accordance with General Instruction B.2. of Form 8-K, the information in this Item 7.01 and Exhibit 99.1 hereto, shall not be deemed "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "***Exchange Act***"), or otherwise subject to the liability of that section, nor shall it be deemed incorporated by reference in any of the Company's filings under the Securities Act of 1933, as amended, or the Exchange Act, whether made before or after the date hereof, regardless of any incorporation language in such a filing, except as expressly set forth by specific reference in such a filing.

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits.

Exhibit No.	Exhibit Description
99.1	Company Presentation.

SIGNATURES

Pursuant to the requirements of the Securities Exchange Act of 1934, the Registrant has duly caused this report to be signed on its behalf by the undersigned hereunto duly authorized.

ACLARIS THERAPEUTICS, INC.

By: /s/ Frank Ruffo
Frank Ruffo
Chief Financial Officer

Date: December 2, 2019

EMPOWERING PATIENTS THROUGH
KINOME INNOVATION

Company Overview

December 2019

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Cautionary Note Regarding Forward-Looking Statements

Any statements contained in this presentation that do not describe historical facts may constitute forward-looking statements as that term is defined in the Private Securities Litigation Reform Act of 1995. These statements may be identified by words such as "believe," "expect," "may," "plan," "potential," "will," and similar expressions, and are based on Aclaris' current beliefs and expectations. These forward-looking statements include expectations regarding Aclaris' development of its drug candidates, including the timing for initiation and completion of clinical trials, the availability of data from these trials and the timing of its regulatory submissions related to these trials. These statements involve risks and uncertainties that could cause actual results to differ materially from those reflected in such statements. Risks and uncertainties that may cause actual results to differ materially include uncertainties inherent in the conduct of clinical trials, Aclaris' reliance on third parties over which it may not always have full control, and other risks and uncertainties that are described in the Risk Factors section of Aclaris' Quarterly Report on Form 10-Q for the quarter ended September 30, 2019, and other filings Aclaris makes with the U.S. Securities and Exchange Commission from time to time. These documents are available under the "SEC filings" section of the Investors page of Aclaris' website at http://www.aclaristx.com. Any forward-looking statements speak only as of the date of this presentation and are based on information available to Aclaris as of the date of this presentation, and Aclaris assumes no obligation to, and does not intend to, update any forward-looking statements, whether as a result of new information, future events or otherwise

This presentation also contains estimates and other statistical data made by independent parties and by us relating to market size and other data about our industry. This data involves a number of assumptions and limitations, and you are cautioned not to give undue weight to such estimates. In addition, projections, assumptions and estimates of our future performance and the future performance of the markets in which we operate are necessarily subject to a high degree of uncertainty and risk.

Strategy: *Development stage biotechnology company focused on immuno-inflammatory diseases*

LEADERSHIP

- Physician/Scientist founded and led
- Kinome experts - combined 300+ years of R&D experience immunology and inflammation
- World class ex-Pfizer kinase and ex-GSK immunology R&D leadership

KINect™ PLATFORM
Proprietary Discovery Engine

- Versatile platform with multiple approaches for difficult to drug kinases in precedented pathways
- Fully integrated discovery and development team
- Dedicated to the design of innovative, kinase targeted medicines for immuno-inflammatory diseases
- Positioning small molecule drug candidates to parallel or exceed efficacy of high value biologics

INNOVATIVE PIPELINE
(investigational drug candidates)

ATI-450
- Oral anti-TNF, anti-IL1, anti-IL6
- Novel target for the potential treatment of various inflammatory indications

ATI-1777-Topical Soft-JAK1/3i
- Innovative treatment limiting systemic exposure for the potential treatment of moderate-to-severe atopic dermatitis (AD)

ATI-2138 - ITK/TXK/JAK3i
- Dual inhibitor of T-cell and cytokine receptor for the potential treatment of auto-immune disease

R&D Leadership Team

Experienced team with deep scientific and operational experience

David Gordon
Chief Medical Officer

Former SVP, R&D at GSK. Led discovery and development teams in Immuno-Inflammation and Dermatology leading to multiple successful NDAs, including NUCALA® & BENLYSTA®

Joseph Monahan, PhD
Exec. VP R&D
(Head of Discovery)

Former Executive Director, Pfizer Inflammation Research and Leader of Global Kinase Technology Team

>95 publications and patents (>30 total on kinases)

Walter Smith
SVP, R&D

Former VP Research & Global Head, Pfizer Inflammation, co-leader of Pfizer Licensing Team

Delivered 8 clinical candidates, 6 INDs and 1 NDA in inflammation and cancer

Jon Jacobsen, PhD
VP, Chemistry

Former Research Fellow and Director, Pfizer Chemistry

>100 publications and patents (15 total on kinases)

Project Lead for PFE JAK Program

Paul Changelian, PhD
VP, Biology

Immunologist/drug discovery leader at pharma (Pfizer & biotech)

Validated JAK 1/3 as target for transplant/RA/psoriasis, leading to approval of XELJANZ®

David R Anderson, PhD
Sr. Director, Discovery, Early Development

Former research project leader at Pfizer. Director of Chemistry at Mnemosyne, Luc, Cadent.

Inventor of 6 clinical candidates and author of 40 peer reviewed publications and patents

Gary DeCrescenzo
SVP, Pharm R&D

Former Exec. Director, Pfizer. Site Head for Medicinal & Structural Chemistry. >100 patents. Co-inventor of multiple drug candidates

Pipeline

Program	Indication(s)	Preclinical	Phase 1	Phase 2	Phase 3
ATI-450 MK2 Inhibitor *Oral*	Rheumatoid Arthritis + Additional Immuno-inflammatory Indication				
ATI-1777 JAK1/JAK3 Inhibitor *Soft Topical*	Atopic Dermatitis (moderate-to-severe)				
ATI-2138 ITK/TXK/JAK3 Inhibitor *Oral*	Psoriasis, Inflammatory Bowel Disease				
JAK1/JAK3 Inhibitor *Oral, gut-restricted*	Inflammatory Bowel Disease				
ITK/JAK3 Inhibitor *Oral, gut-restricted*	Inflammatory Bowel Disease				

The Kinase Opportunity and Challenge

Creating New Medicines Targeting Previously Inaccessible Kinome Targets

Medically Important and Productive Target Class

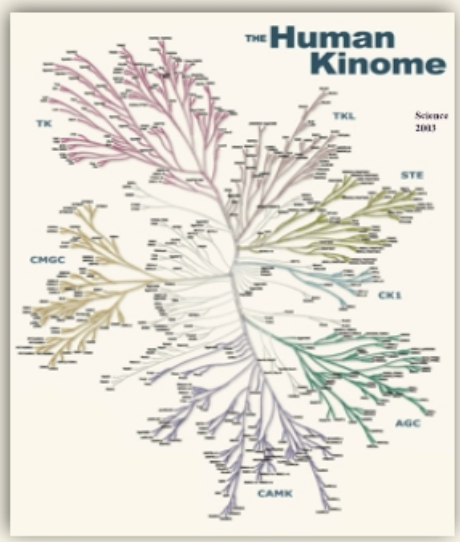
OFEV (nintedanib) capsules 150mg
Bosulif bosutinib
IBRANCE palbociclib
IRESSA
Inlyta
imbruvica (ibrutinib)
ZELBORAF (vemurafenib) tablets
ICLUSIG (ponatinib) tablets
COMETRIQ (cabozantinib)
gleevec (imatinib mesylate) tablets
GILOTRIF (afatinib) tablets
Tarceva erlotinib
ICLUSIG (ponatinib) tablets
Votrient pazopanib tablets 200 mg
Tykerb (lapatinib)
Jakafi ruxolitinib
SPRYCEL dasatinib
Zydelig (idelalisib) tablets
Caprelsa (vandetanib) tablets
Mekinist (trametinib)
XALKORI
Stivarga
SUTENT sunitinib malate
ZYKADIA ceritinib
Nexavar (sorafenib) tablets
LENVIMA (lenvatinib) capsules
Tasigna
XELJANZ (tofacitinib citrate)

~36 Marketed Drugs

~$48B*

Annual Sales of Kinase Drugs

Most Members of the Kinome Remain Unexplored



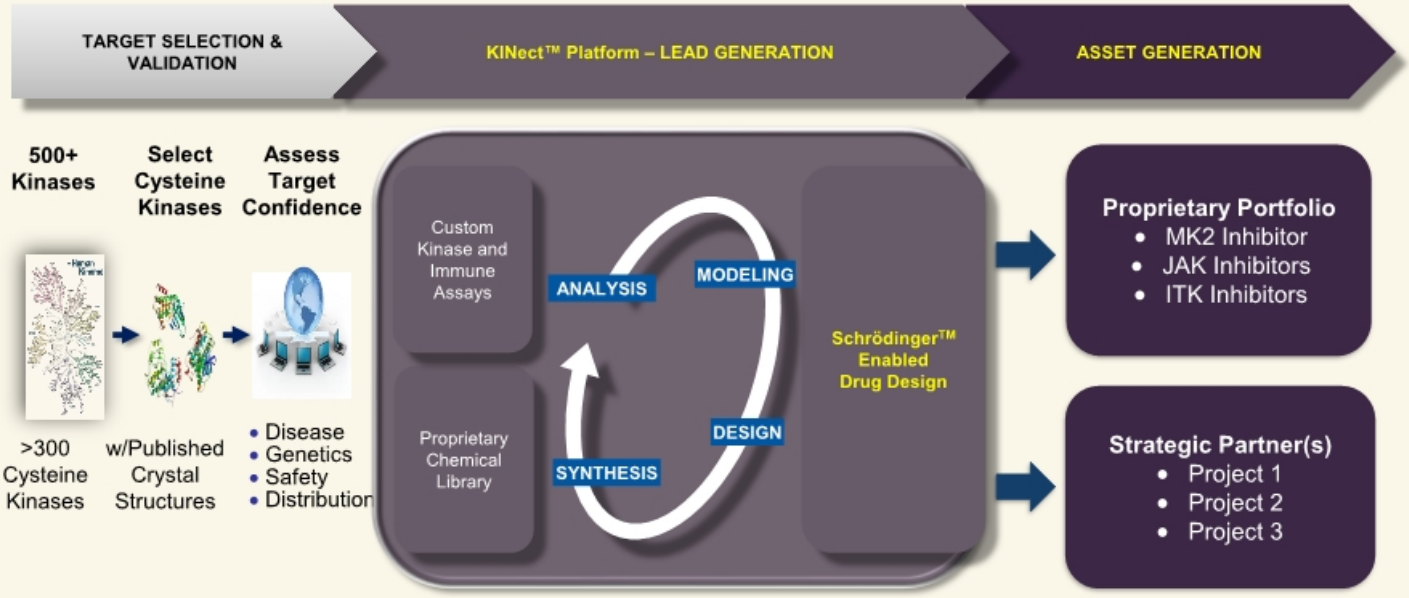
518 Members

>90% of the Human Kinome remains undrugged

These drugs target less then 5% of the kinome

* Bologa C, et al. Unexplored opportunities in the druggable human genome. *Nat Rev Drug Discov*. 2018.

KINect™ Platform
Developing Kinase Drug Candidates Rapidly & Efficiently



- **Forward focus on the cysteinome with use of electrophilic warheads**
- **Leads are chemically tuned**

- **Multiple approaches for difficult to drug kinases**
- **Integrated system:**
  - **Structure-based drug design**
  - **Proprietary library of kinase inhibitor probes**
  - **Layers of validated translatable testing funnels**
- **Faster path to lead optimization**

- **Demonstrated success:**
  - **Substrate selective inhibitors**
  - **Tissue targeted reversible and irreversible kinase inhibitors**
  - **Covalent ITK inhibitors**

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KINect™ Platform Demonstrated Success
Reversible and Covalent

MK2 Inhibitor	Tissue Restricted JAK and ITK Inhibitors	Covalent ITK Inhibitors
Unique Substrate Selective Drug Design	**Tailoring physico-chemical and potency properties**	**Covalent Inhibition: for difficult to target kinase**
• Oral anti-TNF, anti-IL1, and anti-IL6 MK2 kinase inhibitor drug • Novel approach for a difficult to target kinase • ATI-450 (investigational compound) currently in Phase 1 clinical trial	• Potential approaches to achieve efficacy with improved safety • ATI-1777 (investigational compound): Soft, topical drug for the potential treatment of moderate-to-severe AD • Gut-restricted inhibitor for the potential treatment for inflammatory bowel disease	• ITK T cell kinase inhibitors for autoimmune disease • Reversible inhibition largely unsuccessful • Oral and topical covalent drug candidates developed • Oral: ATI-2138 (investigational compound) IND enabling work

Market Overview of Select Inflammatory Indications

	RA (moderate - severe)	Psoriasis (moderate - severe)	Ulcerative Colitis (moderate - severe)	Crohn's (moderate - severe)	Atopic Dermatitis (moderate - severe)
2018E WW Sales[1]	~25B	~15B	~$5B	~$11B	~$1B
Estimated Peak Market (WW)[2]	~$25-30B	~$20-25B	~$8-12B	~$15B	~$8-12B
Prevalent US Moderate/Severe Population[3]	~1,000K+	~1,000-1,300K	~400-500K	~350-450K	~300-700K
Approved Agents (per target)	TNF-alpha: 5	TNF-alpha: 3	TNF-alpha: 2	TNF-alpha: 3	IL-4R: 1
	CD20: 1	IL-12 / IL-23: 2	Integrin α4β7: 1	IL-12 / IL-23: 1	
	JAK: 2	IL-17A: 2	JAK: 1	Integrin α4β7: 1	
	Integrin α4β7: 1	PDE4: 1			
	Other: 3				
Agents in Clinic (per target)	BTK: 9	IL-23: 2	JAK/STAT: 4	JAK/STAT: 5	JAK/STAT: 4
	JAK/STAT: 5	IL-17 / IL17R: 4	IL-23: 4	IL-23: 5	IL-33: 2
	IL-6: 3	JAK/STAT: 2	S1P-R: 2	S1P Receptor: 3	IL-13: 2
	TNF-alpha: 1	Others: 7	Integrins: 2	Integrin α4β7: 1	IL-31: 2
	T-cell Receptor: 1		Others: 12	Others: 12	OX40: 2
	Others: 41				Others: 8
Opportunity for New Treatments	Orals, Improved risk/benefit, novel mechanism	Oral, novel mechanism, improved safety	Gut-restricted (improved safety)	Gut-restricted (Improved safety)	Improved risk/benefit, topical in moderate to severe

* Auster M, et al. Something Big Is Getting Bigger [research note]. New York, NY: Credit Suisse Equity Research; 2019.
1 Estimates of total sales per indication from EvaluatePharma.
2 CS projections: based on US branded pricing.
3 Assumed peak treatable population with biologics/novel agents in the US: RA 350-400k / Psoriasis 300-350k / Ulcerative Colitis 225-275k / Crohn's 225-275k / Atopic Dermatitis 150-200k.

ATI-450: MK2 Inhibitor

(Investigational Drug Candidate)

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MK2 Inhibitor – Potential ORAL Alternative to Injectable, Anti-Cytokine Biologics and JAK Inhibitors for Immuno-Inflammatory Diseases

- MK2* is an attractive drug target because it drives pro-inflammatory cytokine expression
- The effects of inhibiting MK2 mirror the effects of anti-inflammatory biologics[1]
  - **anti-TNF**: HUMIRA®, ENBREL®, REMICADE®
  - **anti-IL1**: KINERET®, ILARIS®, ARCALYST®
  - **anti-IL6**: KEVZARA®, ACTEMRA®
- ATI-450 inhibits MK2 via a novel MOA which involves binding to a drug "pocket" created in the p38α/MK2 complex[2]

* MK2 = Mitogen-activated protein kinase-activated protein kinase 2
** All trademarks are the property of their respective owners.
1 Data on file.
2 Wang C, et al. *J Exp Med*. 2018;215(5):1315-1325.

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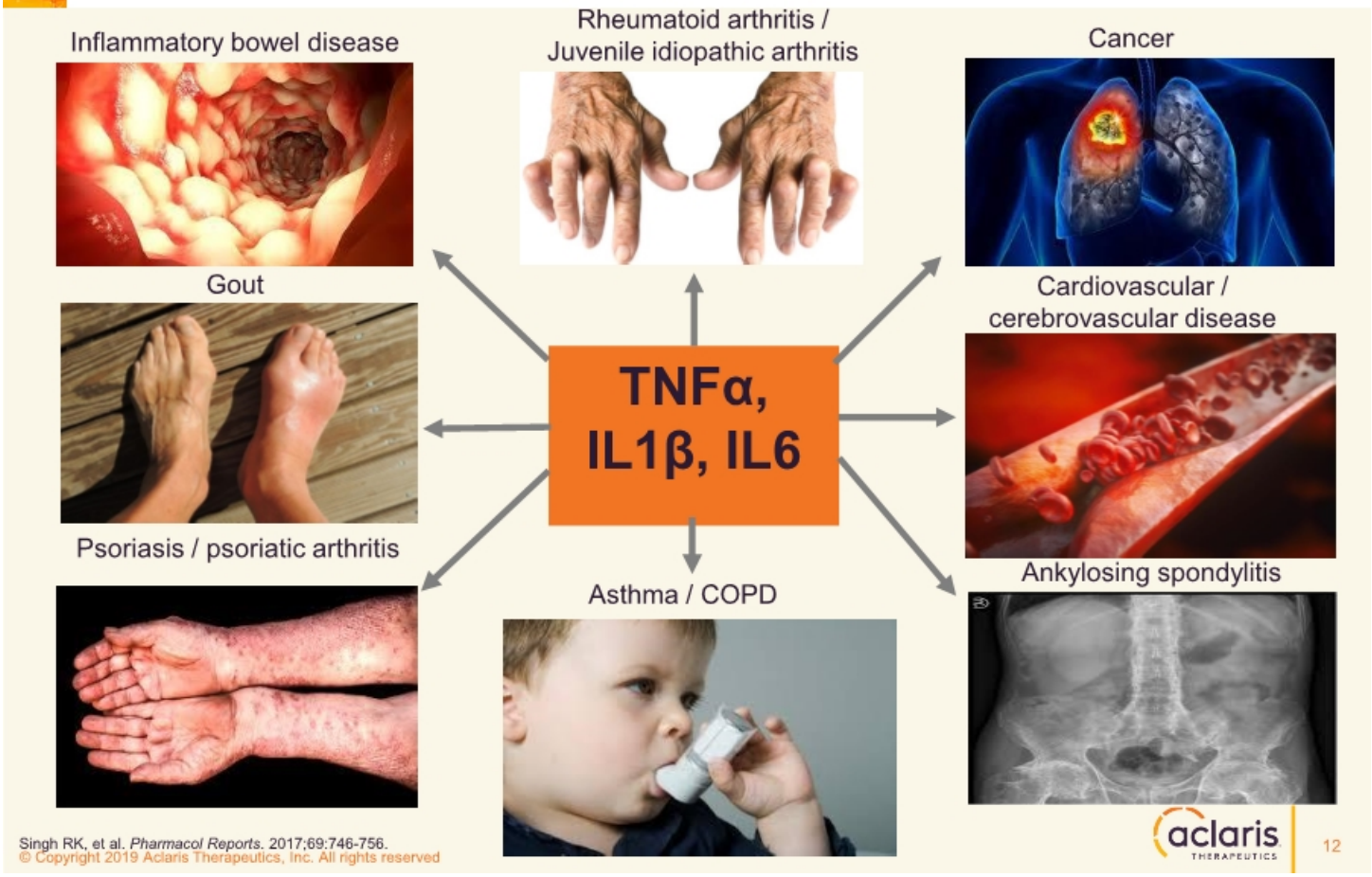
MK2-driven Cytokines are Central to Many Diseases



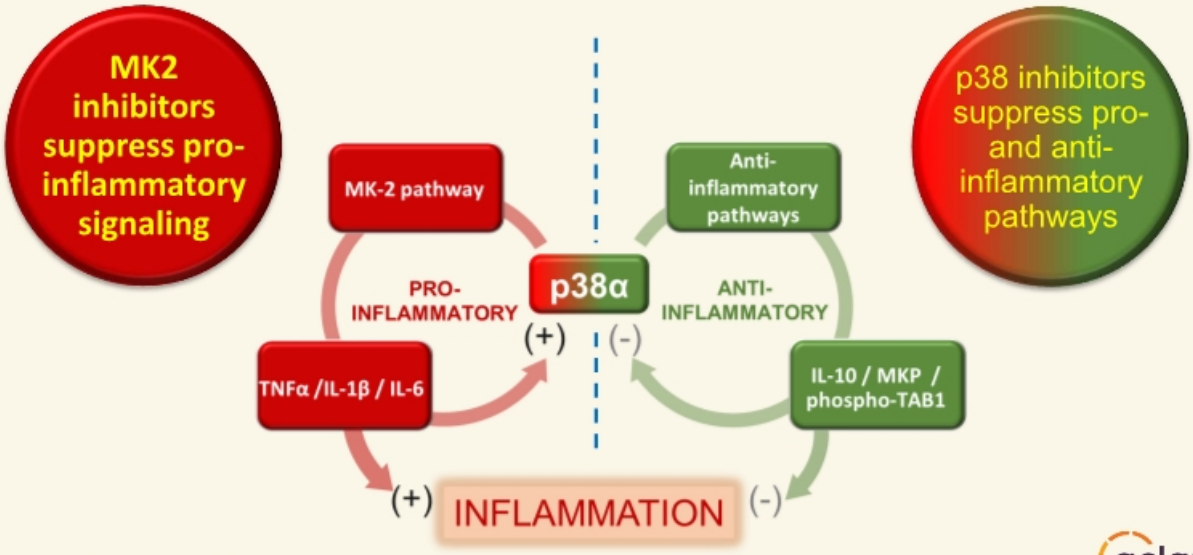
Singh RK, et al. *Pharmacol Reports*. 2017;69:746-756.

Evolution in Understanding of a Well-Known Path

The Path From p38α to MK2

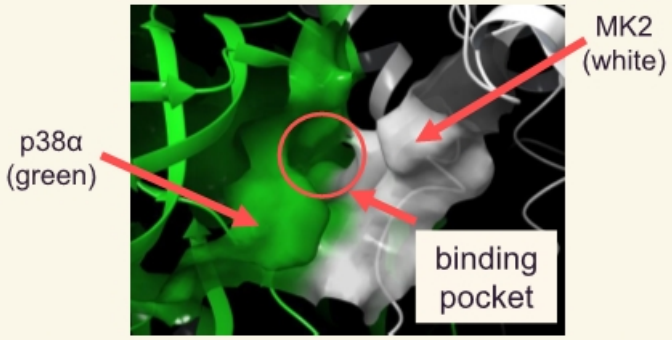
p38α was the original therapeutic target for suppressing TNFα and other pro-inflammatory cytokines

- However, first generation p38α inhibitors were not selective resulting in multiple safety issues including liver, kidney, GI, and skin toxicity.
- Second generation p38α inhibitors demonstrated a lack of sustained activity in certain diseases such as RA and IBD and could not completely overcome toxicity.
- MK2 inhibitors became recognized as a more selective and targeted approach to this path.

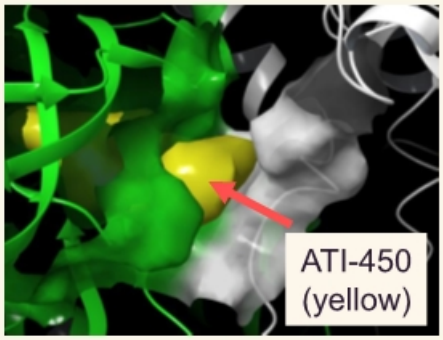


* Schindler J, et al. *J Dent Res.* 2007;86(9):800-811.

Capturing MK2 in an Inactive State



Crystal structure of the p38α/MK2 complex



ATI-450 (yellow) docked in the pocket

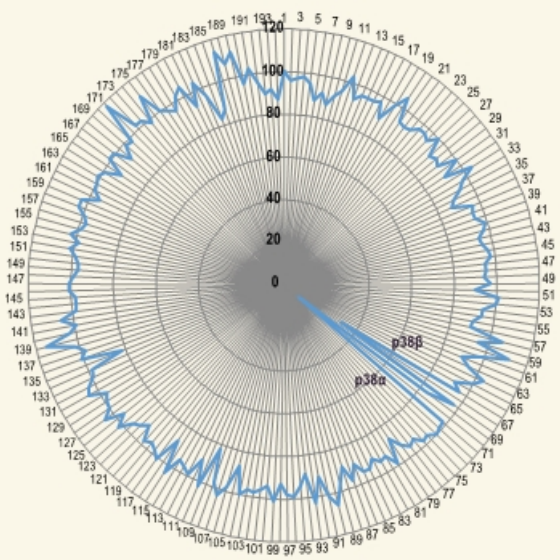
- In the nucleus, inactive MK2 and p38α dock in a high affinity complex that exhibits a binding pocket formed by juxtaposed walls of both proteins
- Aclaris MK2 inhibitors bind to both walls of the pocket, stabilizing the complex and preventing MK2 activation

Aclaris MK2 inhibitors lock MK2 in a catalytically inactive state – a unique MOA

* Wang C, et al. *J Exp Med*. 2018;215(5):1315-1325.

ATI-450 selectivity

Human Kinome Selectivity[1]



- ATI-450 (5μM) was tested vs 193 kinases
- >350-fold binding selectivity on all kinases in this panel except p38α and p38β

MK2 Pathway Selectivity

ATI-450 is highly selective for the p38/MK2 complex vs. other p38 substrates[1]

Assay	Fold Selective
p38α/MK2	1
p38α/ATF2	700
p38α/PRAK	750

ATI-450 binds to the p38α/MK2 complex with higher affinity than either p38 or MK2 alone**

Assay	Fold Selective
p38α/MK2	1
p38α/p38tide*	51
MK2/HSP27	>550

* Optimized p38 peptide substrate
** Data on file.
1 Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.

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Animal Models Supporting the Development of ATI-450 in Inflammatory Diseases

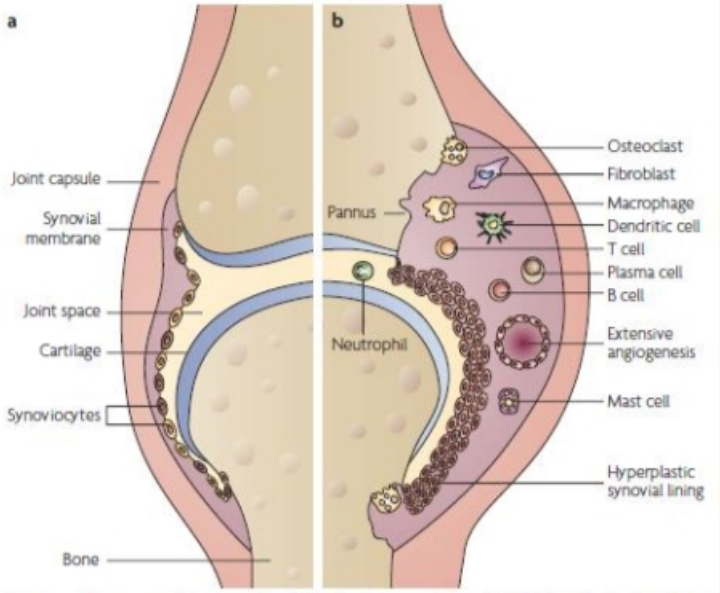
Therapeutic Area	Animal Model	Reference
Rheumatoid Arthritis / Psoriatic Arthritis	Rat streptococcal cell wall arthritis model • Protection against bone deterioration • Protection against lethality Inhibition of cellular IL1β mRNA stability & translation	Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.
Inflammatory Bowel Disease	Adoptive transfer mouse model of colitis • Endoscopy scores show disease control • Decreased inflammatory infiltrate • Protected structural integrity of mucosa	Strasser S, et al. *Integrative Biology*. 2019;11(7):301-314.
Cryopyrin-Associated Periodic Syndromes (CAPS)	Murine NOMID (severe form of CAPS) transgenic model Human CAPS PBMC* IL1β modulation	Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.

* PBMC = Peripheral blood mononuclear cells

ATI-450 Blocks MK2 – Potential Effect in Rheumatoid Arthritis

MK2 is a key regulator of essential pathogenic signals in chronic inflammatory and autoimmune diseases

Normal Joint | RA Joint



Strand V, et al. *Nat Rev Drug Discov*. 2007;6(Jan 2007):75-92.

Cells	Cytokines
Monocyte	**TNFα**
Macrophage	**IL1β**
Osteoclast	**IL1α**
Epithelial Cells	**IL6**
RA Syn Fibroblast	**IL18**
Chondrocytes	**RANKL**

ATI-450: for bold items above data on file and Wang C, et al. *J Exp Med*. 2018;215 (5):1315-1325.

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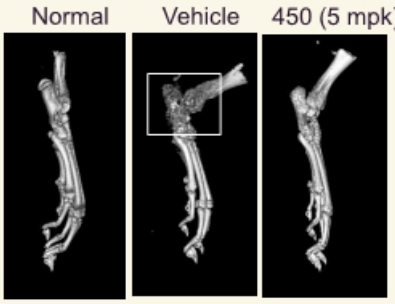
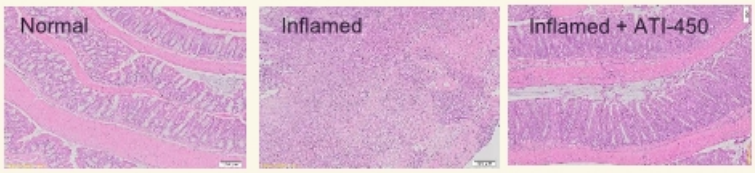
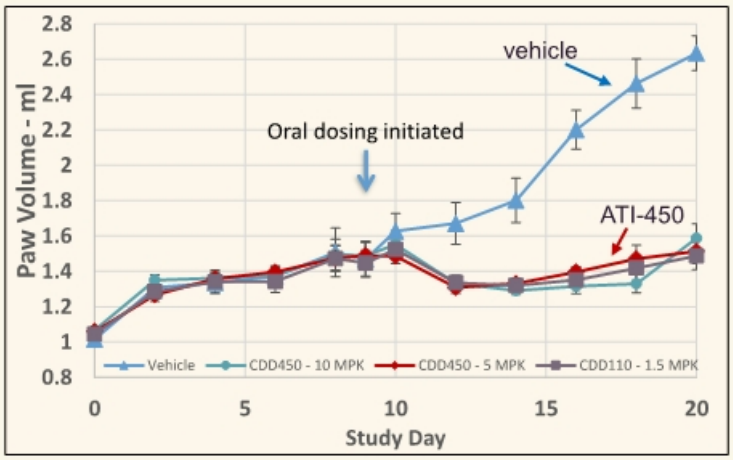
In vivo Preclinical Data of MK2 Pathway Inhibitor ATI-450

Joint Protection in Rat Arthritis Model[1]

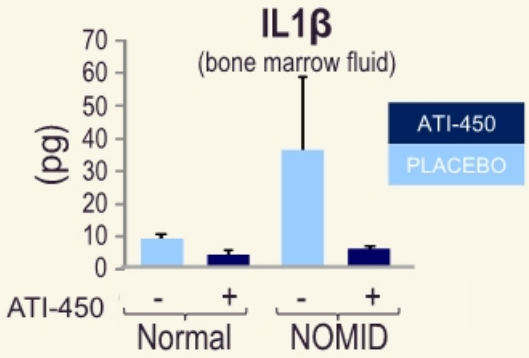


Blockade of Gut Inflammatory Infiltrate in Murine Adoptive Transfer Ulcerative Colitis Model[2]





Cytokine Modulation in Orphan Autoinflammatory Disease (CAPS)[1]



1 Wang C, et al. *J Exp Med*. 2018;215(5):1315-1325.
2 Strasser S, et al. *Integrative Biology*. 2019;11(7):301-314.

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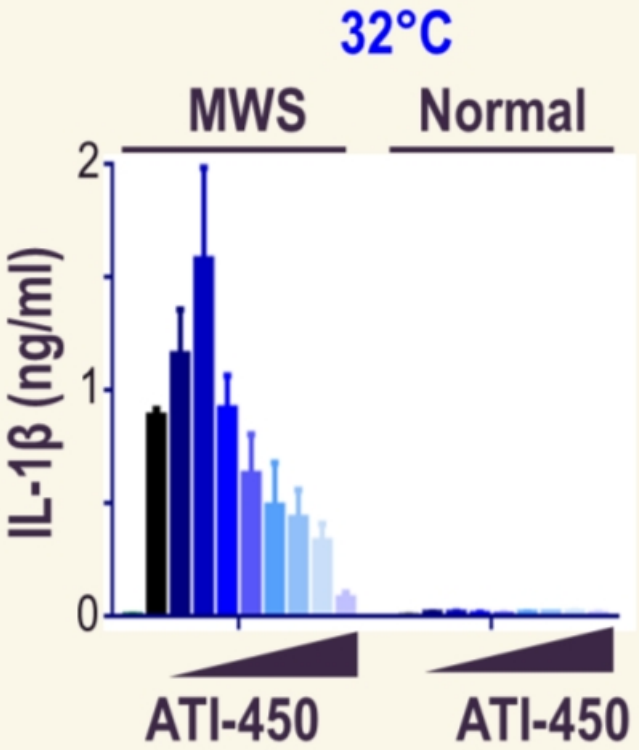
Ex vivo Preclinical Data: ATI-450 Inhibits IL1β Expression in PBMCs from a CAPS Patient



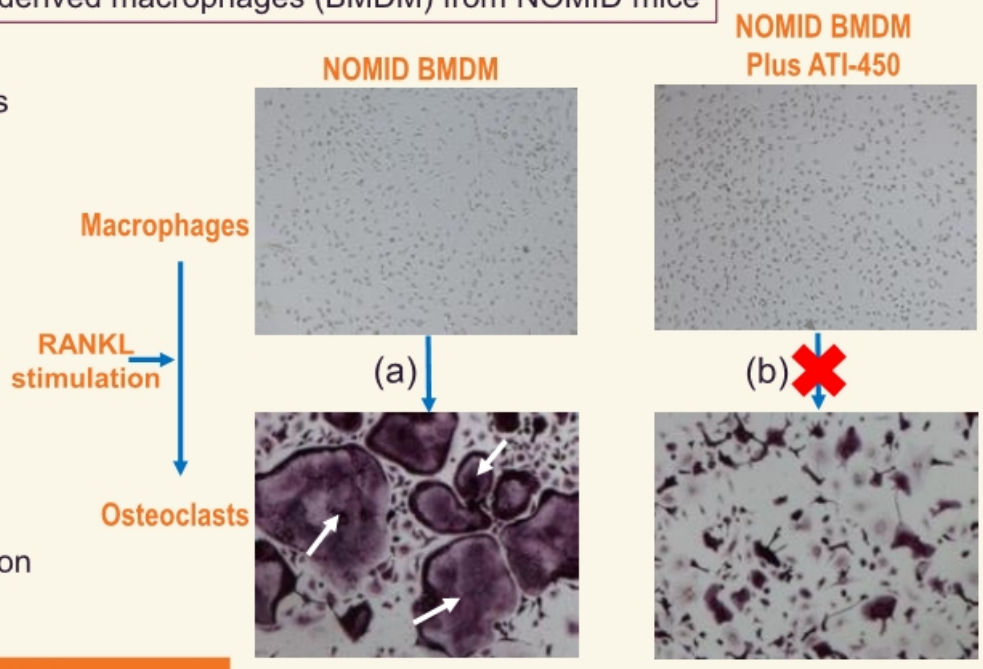
- Peripheral blood mononuclear cells (PMBCs) were isolated from CAPS patients and healthy controls.
- In CAPS patients (Muckle Wells Syndrome; MWS), IL1β expression is triggered by exposure to low temperatures.
- CAPS PBMCs spontaneously produced high amounts of IL1β at 32°C but not at 37°C.
- ATI-450 blocks temperature stress induced IL1β production.

* Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.

Mouse Model: ATI-450 Inhibits RANKL-induced Osteoclastogenesis

Bone marrow derived macrophages (BMDM) from NOMID mice

- In CAPS, osteoclastogenesis gives rise to low bone mass (osteopenia)
- (a) When bone marrow derived macrophages (BMDM) from NOMID mice are stimulated with RANKL (RANK ligand), they differentiate into osteoclasts
- (b) ATI-450 blocks this macrophage differentiation



ATI-450 inhibits RANKL-stimulated macrophage differentiation into osteoclasts from NOMID mice

* Wang C, et al. *J Exp Med*. 2018;215(5):1315-1325.

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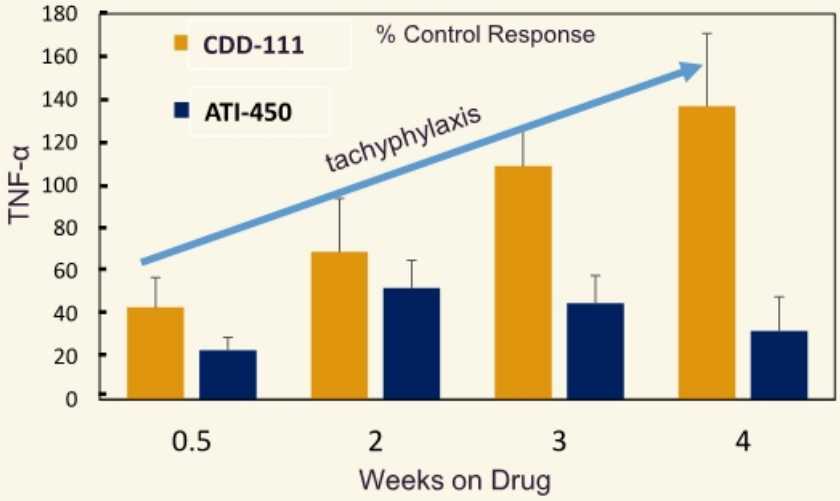
Mouse Model: LPS-Induced TNFα Production

ATI-450 demonstrated durable response (no tachyphylaxis)

- Global p38 inhibitor CDD-111 lost inhibition over time
- **MK2 inhibitor ATI-450 (investigational compound) demonstrated durable response (no tachyphylaxis)**



- Conventional p38 (CDD-111) and MK2i (ATI-450) administered to mice in feed starting day 1 and continuing through day 28
- At the time point indicated, mice were LPS challenged and blood TNFα levels determined

* Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.

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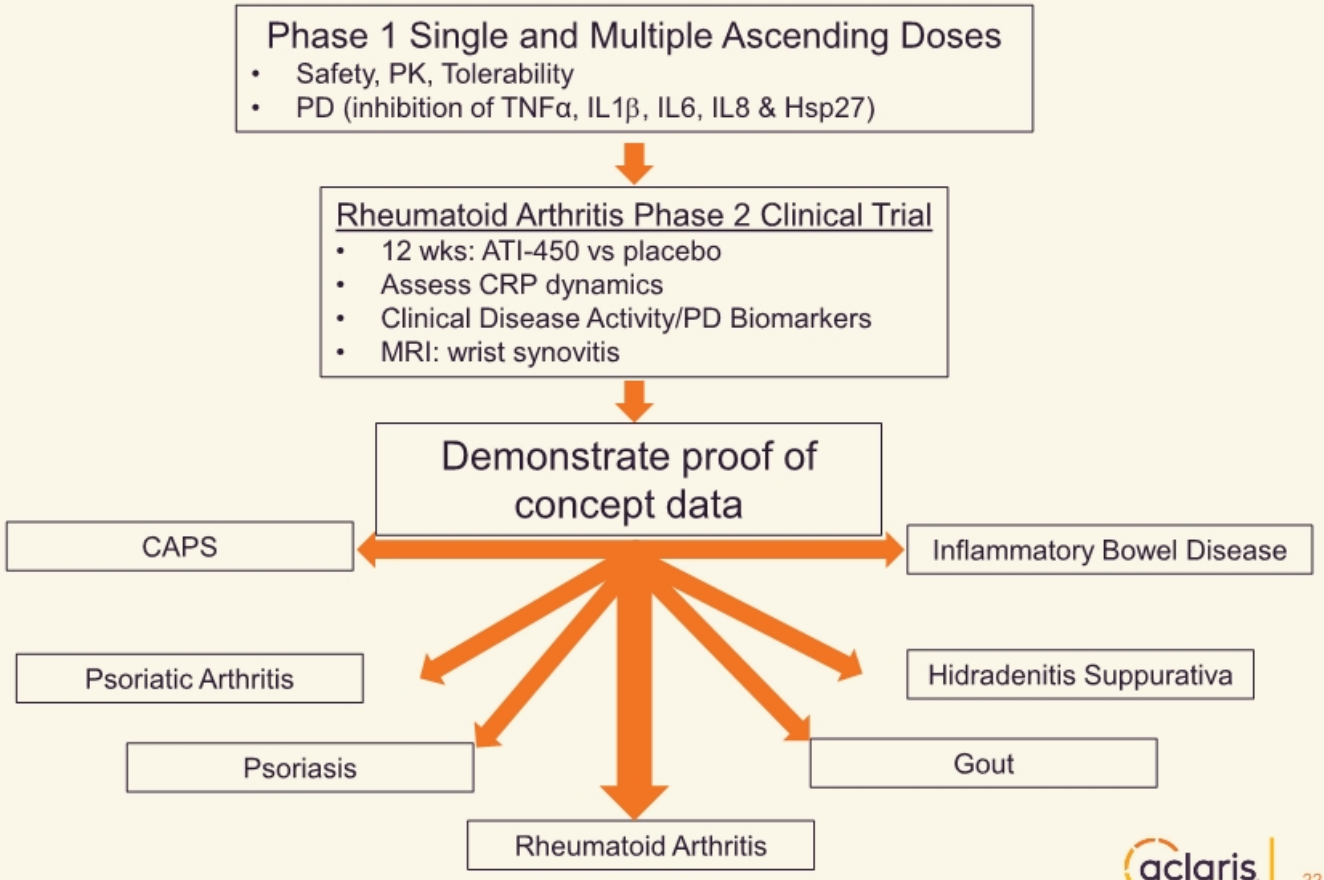
ATI-450 Clinical Development



MK2 inhibitor ATI-450 Summary

- Well-known pathway
- Discovered an approach to drug the target
- Novel mechanism designed to block inflammation
  - ✓ Multiple inflammatory cytokines impacted
  - ✓ Key RA inflammatory cell types impacted
  - ✓ Lock MK2 in a catalytically inactive state – a unique MOA
  - ✓ Broad IP issued
- Oral option for numerous diseases currently treated by biologics
  - ✓ Robust efficacy in a range of inflammation and mouse cancer models[1,2]
- Phase 1 trial underway
- Proof of concept Phase 2 trial in RA expected to begin first half 2020
  - ✓ To demonstrate clear pharmacodynamic effect and no tachyphylaxis
  - ✓ To demonstrate early signs of efficacy in a well understood disease
- Other inflammatory indications under consideration

1 Murali B, et al. *Cancer Res.* 2018;78(19):1-13.
2 Wang C, et al. *J Exp Med.* 2018;215(5):1315-1325.

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ATI-1777 (Topical Soft-JAK Inhibitor)

(Investigational Drug Candidate)

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ATI-1777 (Topical Soft-JAK Inhibitor)

Novel approach for moderate to severe Atopic Dermatitis

- Atopic dermatitis (AD) is a disease of unknown origin that usually starts in early infancy and is typified by pruritus, eczematous lesions, xerosis (dry skin), and lichenification on the skin (thickening of the skin and increase in skin markings).[1]
  - ✓ Large and growing market – Projected to be $8-12 billion at peak (moderate-to-severe AD)[2]
  - ✓ Unmet need for effective and safe topical treatment for AD
  - ✓ Systemic and topical JAK inhibition has demonstrated promising results in clinical trials for treating pruritus and inflammation in AD[3]
  - ✓ In AD, a compromised skin barrier means that a topically dosed JAK inhibitor might result in pharmacologically active systemic drug levels
- Topical soft-JAK inhibitor has potential to achieve efficacy with improved safety
  - ✓ Achieve efficacy in skin while minimizing systemic JAK inhibitor toxicity
  - ✓ JAK1/3 selectivity minimizes JAK2 toxicities given compromised skin barrier
- Topical formulations being optimized into a differentiated, patient-friendly emollient formulation (topical spray vs cream/ointment)
- First in human studies planned for second half 2020 in moderate-to-severe AD

1 https://emedicine.medscape.com/article/1049085-overview. Last accessed 11-1-19.
2 Auster M, et al. Something Big Is Getting Bigger [research note]. *Credit Suisse Equity Research*; 2019.
3 Shreberk-Hassidim R, et al. *J Am Acad Dermatol*. 2017;Apr;76(4):745-753.

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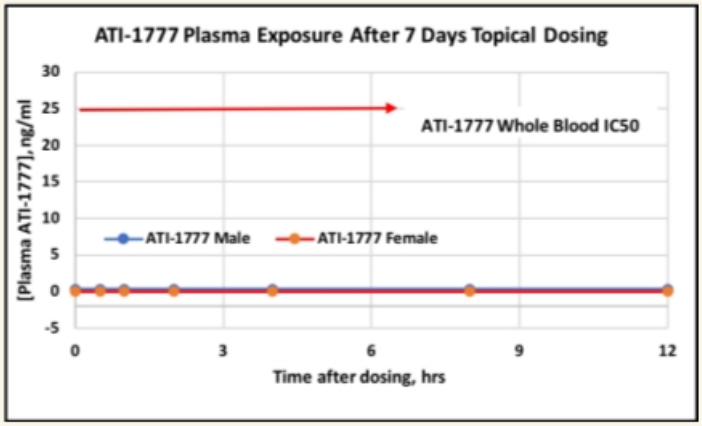
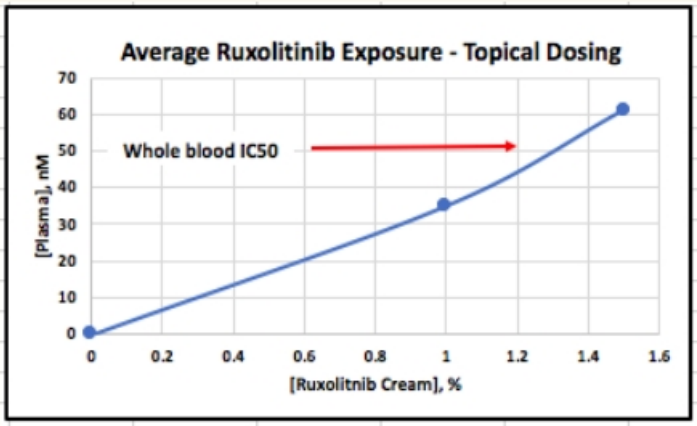
Minipig Model: ATI-1777 Nonclinical Safety Program TK Data

Tolerability/Toxicokinetic with 7-day dermal administration (non-GLP)

- No adverse effects noted (10% body surface area, QD)
- Bleeds at 0.5, 1, 2, 4, 8, 12, and 24 hours post-application: Days 1 and 6
- All plasma samples were below limit of quantification (<0.50 ng/mL) – well below cellular IC_{50}



MINIPIG[1]



HUMAN[2,3]

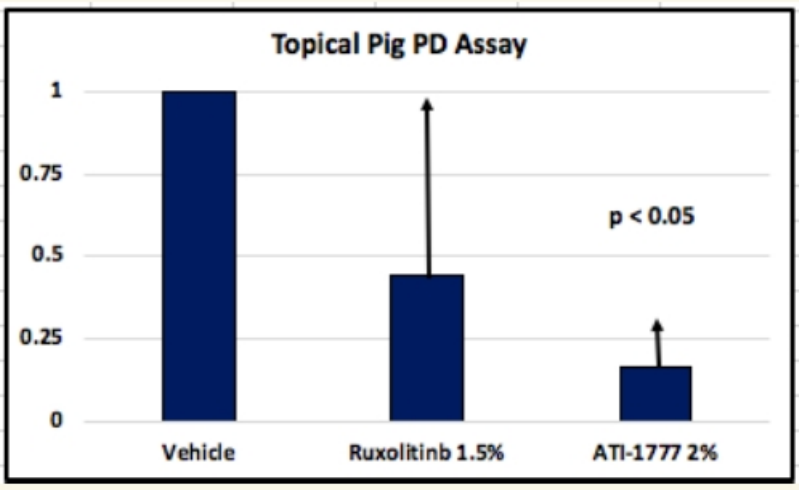
1 Data on file.
2 Chen X, et al. *Clin Pharmacol Drug Dev*. 2013;3(1):34–42.
3 Punwani N, et al. *Br J Dermatol*. 2015;173:989–997.

Porcine Model: ATI-1777 Blocks IL15 Induced CCL8 mRNA

Apply formulation to back of pig, wait 1 hr

Intra-dermal Injection of porcine IL15, wait 3 hr

Harvest 6 mm biopsy, prepare RNA, measure CCL8 by qPCR



- Single application of 2% ATI-1777 development formulation significantly inhibits IL15 induced gene induction (CCL8).
- Clinical topical formulation of 1.5% ruxolitinib does not significantly inhibit IL15 (CCL8) induction.

* Data on file

ATI-1777: Topical Soft-JAK Inhibitor to Target Moderate-to-Severe AD

Formulate a topical atopic dermatitis therapy which meets the medical, aesthetic and compliance needs of patients and physicians

Approach

- Designed to be:
  - "Soft" drug to minimize the potential for systemic immunosuppression
  - JAK1/3 selective to minimize JAK2 inhibition toxicity
  - Delivered in a patient-friendly formulation to clearly differentiate it from other topical therapies

Status

- Plan to study in patients with moderate-to-severe AD
- IND-enabling preclinical safety program initiated
- Next key milestone: First In Human - 2H2020

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ATI-2138 (ITK/TXK/JAK3)

(Investigational Drug Candidate)

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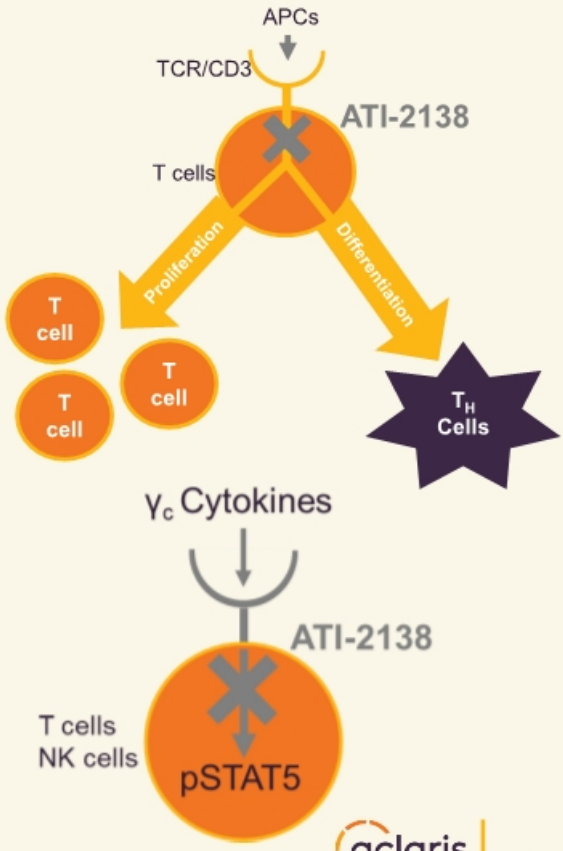
ATI-2138: Covalent ITK/TXK/JAK3 Inhibitor

- ATI-2138 covalently blocks ITK/TXK/JAK3*
  - ✓ ITK/TXK required for T-cell receptor signaling
  - ✓ JAK3 required for γc cytokines (IL-2/4/7/9/15/21)
  - ✓ Targeting both with a single drug may produce synergistic efficacy - expected to have few off-target effects
  - ✓ PD effects persist after plasma clearance
  - ✓ Efficacy in rat arthritis and mouse colitis
- All T-cell mediated autoimmune diseases are potentially targetable
- Next milestones
  - ✓ IND submission in 4Q20/1Q21
  - ✓ First In Human - 1H 2021



* Data on file

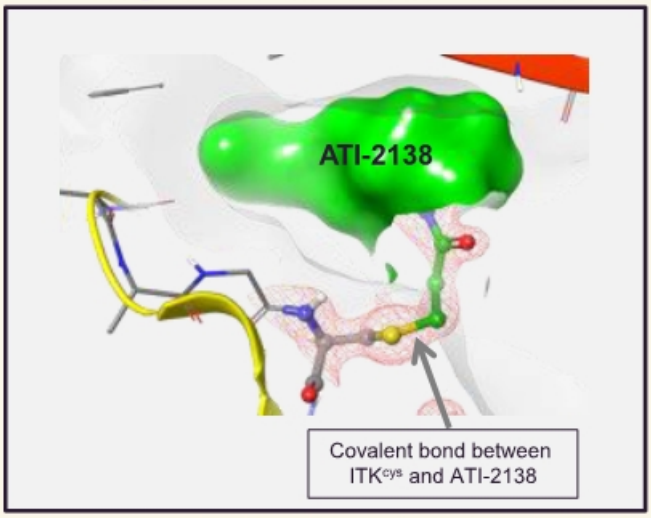
ATI-2138 is a Potent Covalent Inhibitor

Cellular Inhibition of JAK and ITK/TXK

Assay Description	ATI-2138 IC_{50} (nM)	Assay
ITK/TXK activity	7	Jurkat pPLCγ-1
JAK1/3 activity	20	PBMC pSTAT-5
BTK activity	52	Ramos pPLCγ-2
Both ITK/TXK and JAK3	13	HWB αCD3/IL15 IFNγ

ATI-2138 potently inhibits ITK/TXK and JAK3 in cells and in whole blood

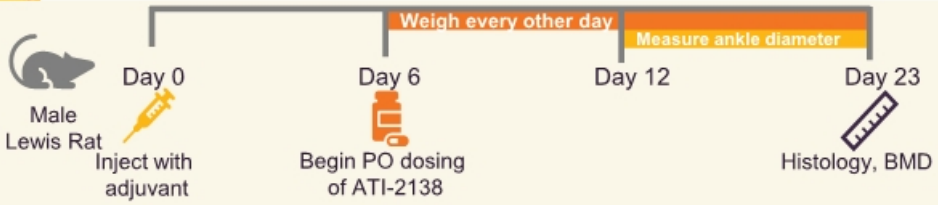
Co-Crystal Structure of ATI-2138/ITK



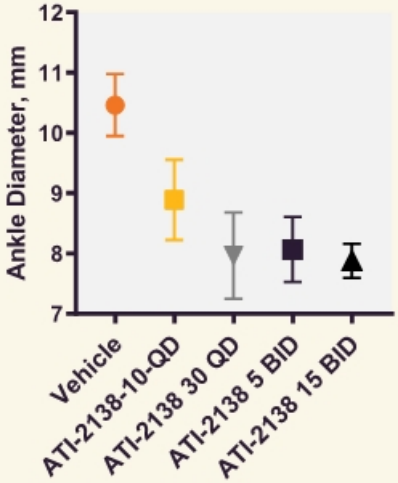
Crystal structure definitively shows ATI-2138 covalent binding to ITK

* Data on file

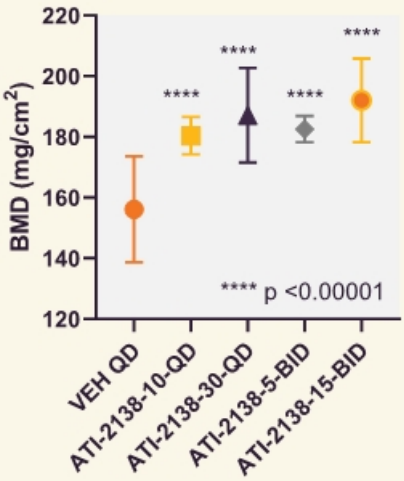
Rat Adjuvant Induced Arthritis (AIA) Model:
ATI-2138 Reduced Inflammation and Protected Bone

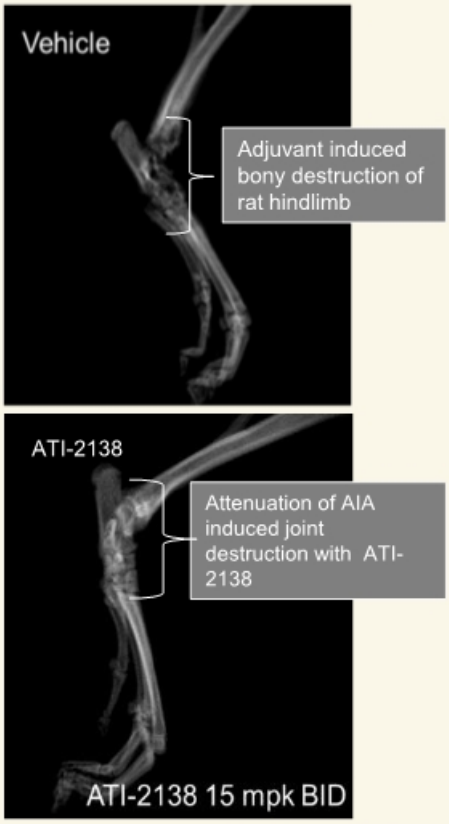


Day 23 ankle diameter



ATI-2138 bone mineral density in rat AIA study





ATI-2138 reduced inflammation and bone mineral density loss

* Data on file

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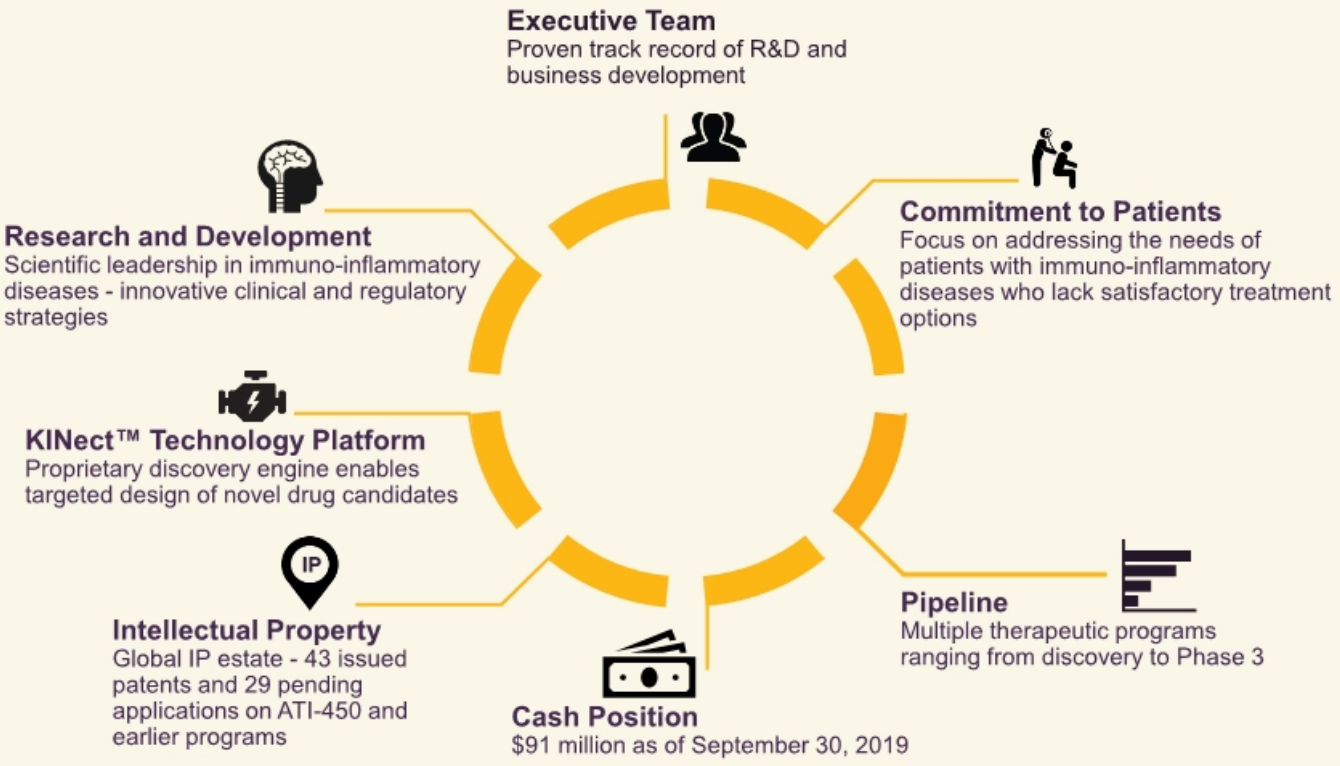
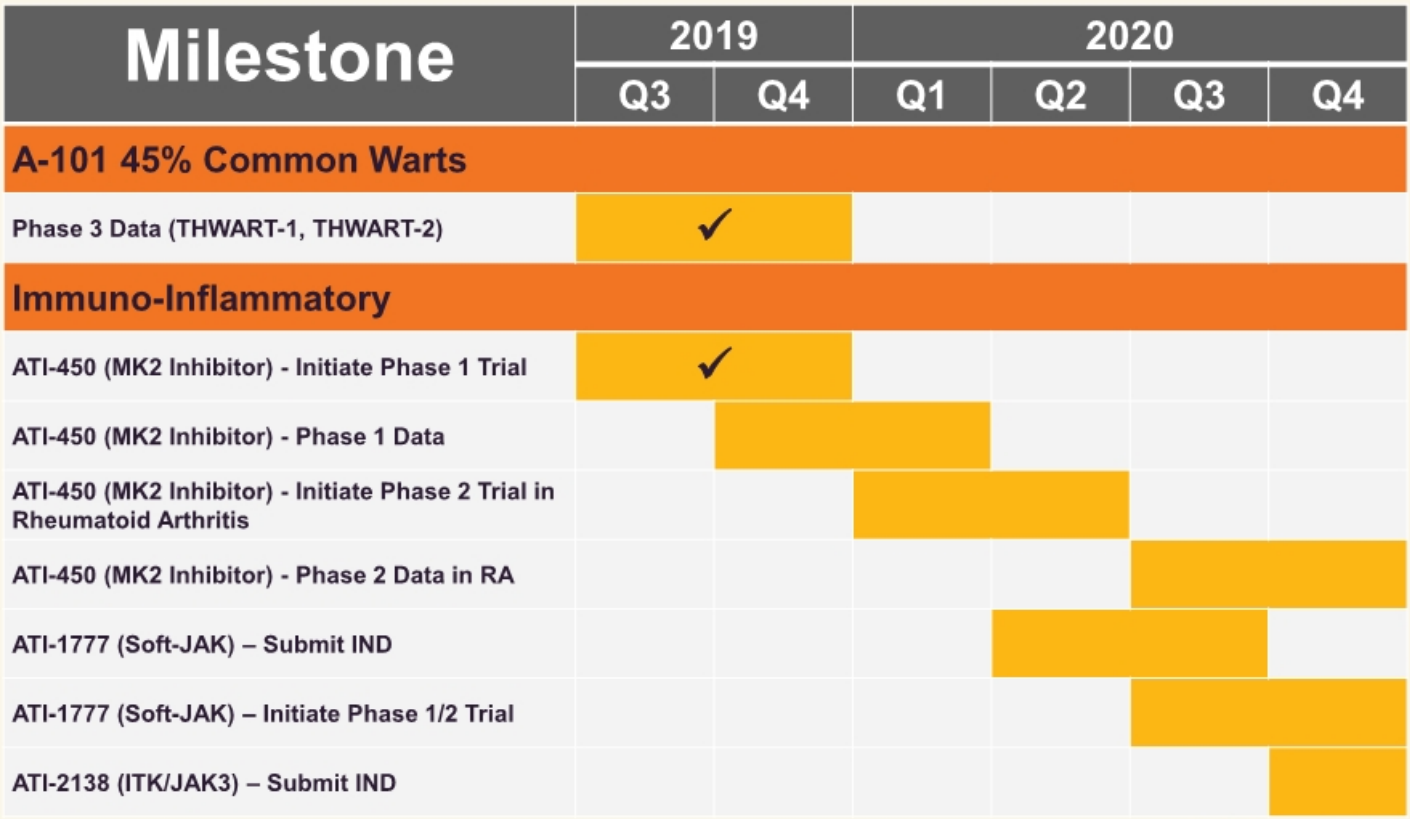
Biopharmaceutical Company



Catalysts



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KINOME INNOVATION

THANK YOU

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