UNITED STATES SECURITIES AND EXCHANGE COMMISSION

Washington, D.C. 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):
March 15, 2022

Ventyx Biosciences, Inc.

(Exact name of registrant as specified in its charter)

Delaware (State or other jurisdiction of incorporation) 001-40928 (Commission File Number) 83-2996852 (IRS Employer Identification No.)

662 Encinitas Blvd., Suite 250
Encinitas, CA 92024
(Address of principal executive offices, including zip code)

(760) 593-4832 (Registrant's telephone number, including area code)

Not Applicable

(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 (see General Instruction A.2. below):

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:

Trading Symbol(s)

Common Stock, \$0.0001 par value per share

Name of exchange on which registered

VTYX

The Nasdaq Global Select Market

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 $\ oxtimes$

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. \Box

Item 7.01 Regulation FD Disclosure.

On March 16, 2022, representatives of Ventyx Biosciences, Inc. ("Ventyx") will be attending meetings with investors, analysts and others in connection with Barclays Global Healthcare Conference taking place on March 16, 2022 in Miami, Florida. During these meetings, Ventyx will present the slides attached as Exhibit 99.1 to this Current Report on Form 8-K, which are incorporated herein by reference.

In accordance with General Instruction B.2. of Form 8-K, all of the information furnished in this Item 7.01 and Item 9.01 (including Exhibit 99.1) shall not be deemed to be "filed" for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the "Exchange Act"), and shall not be incorporated by reference in any filing under the Securities Act of 1933, as amended, or in any filing under the Exchange Act, except as shall be expressly set forth by specific reference in such a filing.

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits.

Exhibit No. Description

99.1 <u>Corporate Presentation, dated March 15, 2022</u>

104 Cover Page Interactive Data File (embedded within the Inline XBRL document).

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.

VENTYX BIOSCIENCES, INC.

By: /s/ Raju Mohan

Raju Mohan, Ph.D. Chief Executive Officer

Date: March 15, 2022



Forward Looking Statements

Ventyx cautions you that statements contained in this presentation regarding matters that are not historical facts are forward-looking statements. These statements are based on Ventyx's current beliefs and expectations. Such forward-looking statements include, but are not limited to, statements regarding: management's belief that three of Ventyx's product candidates are potentially best-in-class; the anticipated timing of commencement, enrollment and completion of clinical trials for Ventyx's product candidates; the anticipated timing of releasing data for the VTX958 MAD trial and advancing VTX958 into Phase 2 trials in psoriasis, psoriatic arthritis and Crohn's disease; the anticipated timing for releasing top-line data for the Phase 2 randomized, placebo-controlled clinical trial for VTX002 and the expectation that such trial, along with an additional Phase 3 trial, may serve as the first of two pivotal trials required for registration; the potential of Ventyx's product candidates to address a broad range of immune-mediated diseases; the anticipated timing for reporting data from the Phase 1 trial for VTX2735 in healthy volunteers and plans for advancing VTX2735 into one or more proof-of-concept trials; anticipated timing for submitting an IND application for VTX3232; plans to advance Ventyx's product candidates; and the expected timeframe for funding Ventyx's operating plan with current cash, cash equivalents and marketable securities. The inclusion of forward-looking statements should not be regarded as a representation by Ventyx that any of its plans will be achieved. Actual results may differ from those set forth in this presentation due to the risks and uncertainties inherent in Ventyx's business, including, without limitation: potential delays in the commencement, enrollment and completion of clinical trials; disruption to our operations from the ongoing global outbreak of the COVID-19 pandemic, or from the ongoing military conflict in Ukraine and the imposition of sanctions against Russia and Belarus, including clinical trial delays; Ventyx's dependence on third parties in connection with product manufacturing, research and preclinical and clinical testing; the results of preclinical studies and early clinical trials are not necessarily predictive of future results; the success of Ventyx's clinical trials and preclinical studies for its product candidates; interim results do not necessarily predict final results and one or more of the outcomes may materially change as the trial continues and more patient data become available and following more comprehensive audit and verification procedures; regulatory developments in the United States and foreign countries; unexpected adverse side effects or inadequate efficacy of our product candidates that may limit their development, regulatory approval and/or commercialization, or may result in recalls or product liability claims; Ventyx's ability to obtain and maintain intellectual property protection for its product candidates; Ventyx may use its capital resources sooner than it expects; and other risks described in Ventyx's prior communications and Ventyx's filings with the Securities and Exchange Commission (SEC), including under the heading "Risk Factors" in Ventyx's Annual Report on Form 10-Q filed on November 18, 2021, and any subsequent filings with the SEC. You are cautioned not to place undue reliance on these forward-looking statements, which speak only as of the date hereof, and Ventyx undertakes no obligation to update such statements to reflect events that occur or circumstances that exist after the date hereof. All forward-looking statements are qualified in their entirety by this cautionary statement, which is made under the safe harbor provisions of the Private Securities Litigation Reform Act of

Neither the SEC nor any state securities commission has approved of the securities of the Company or passed upon the accuracy or adequacy of this presentation. Any representation to the contrary is a criminal offense. Except as otherwise indicated, this presentation speaks as of the date hereof.

This presentation includes statistical and other industry and market data that we obtained from industry publications and research, surveys and studies conducted by third parties as well as our own estimates of potential market opportunities. Industry publications and third-party research, surveys and studies generally indicate that their information has been obtained from sources believed to be reliable, although they do not guarantee the accuracy or completeness of such information. Our estimates of the potential market opportunities for our products include several key assumptions based on our industry knowledge, industry publications, third-party research and other surveys, which may be based on a small sample size and may fail to accurately reflect market opportunities. While we believe that our internal assumptions are reliable, such assumptions have not been verified by any third party. The industry in which we operate is subject to a high degree of uncertainty and risk due to a variety of important factors that could cause results to differ materially from those expressed in the estimates made by third parties and by us.

Trademarks in this presentation are the property of their respective owners and used for informational and education purposes only





INTRODUCTION & PIPELINE

VTX958 | TYK2 Inhibitor | Phase 1

VTX002 | S1P1R Modulator | Phase 2

VTX2735 | Peripheral NLRP3 Inhibitor | Phase 1

CNS NLRP3 Inhibitor | Pre-clinical

Summary | Milestones & highlights

Our Leadership Team

MANAGEMENT



Raju Mohan, PhD
CHIEF EXECUTIVE OFFICER,
FOUNDER



Martin Auster, MD CHIEF FINANCIAL OFFICER



Chris Krueger, JD
CHIEF BUSINESS OFFICER



John Nuss, PhD
CHIEF SCIENTIFIC OFFICER



Jörn Drappa, MD, PhD CHIEF MEDICAL OFFICER

BOARD OF DIRECTORS

Sheila Gujrathi, MD EXECUTIVE CHAIR, VENTYX

Aaron Royston, MD MANAGING PARTNER, VENBIO Jigar Choksey
PRINCIPAL, THIRD POINT

Somu Subramaniam MANAGING PARTNER, NEW SCIENCE VENTURES Richard Gaster, MD, PhD MANAGING PARTNER, VENBIO

William White
CHIEF FINANCIAL OFFICER, AKERO
THERAPEUTICS

Raju Mohan, PhD
CHIEF EXECUTIVE OFFICER, VENTYX



Our Mission: To become a Leading Immunology Company

Underpinned by Strong Drug Discovery and Development Capabilities

WITH THREE, DIFFERENTIATED, CLINICAL-STAGE CANDIDATES

and a deep pipeline of preclinical programs that target immune-mediated and inflammatory disease indications

OUR INTERNALLY-DISCOVERED SMALL MOLECULE DRUGS

allow us to own 100% commercial rights to our entire portfolio with long patent lives for all product candidates

OUR EXPERIENCED TEAM AND OUR INTERNAL R&D ENGINE

continue to generate candidates with potential to address diseases with high unmet need



Broad Pipeline of Candidates With Multiple Near-Term Catalysts

Addressing Established Inflammatory and Immunology Markets with a Wholly Owned Product Portfolio

TARGET	PROGRAM	PRECLINICAL	PHASE 1	PHASE 2	PHASE 3	NEXT ANTICIPATED MILESTONES
TYK2	VTX958	Potential indications inclu	ude psoriasis, psoriatic ar	thritis, Crohn's disease and oth	ners	Complete Phase 1 MAD H1 2022 Initiate Phase 2 POC trial(s) H2 2022
S1P1R	VTX002	Ulcerative Colitis				Report topline Phase 2 data 2023
NLRP3 Peripheral	VTX2735	Potential indications inclu	ude cardiovascular, hepat	ic, renal, and rheumatologic dis	seases	Complete Phase 1 H1 2022 Initiate Phase 2 POC trial(s) H2 2022
NLRP3 CNS-penetrant	VTX3232	Neuroinflammatory disea	ases			Complete IND-enabling studies 2022 File IND Q4 2022



Pipeline Targeting Large Well-Established Markets

INDICATION*	PATIENTS IN THE U.S.	GLOBAL DRUG REVENUE* (2020)	TARGET POPULATION
Psoriasis Dermatology	~8M	~\$20B	25-30% MODERATE-TO-SEVERE
Crohn's disease IBD	~700K	~\$13B	30-40% MODERATE-TO-SEVERE
Ulcerative colitis IBD	~1M	~\$7B	30-40% MODERATE-TO-SEVERE
Psoriatic arthritis Rheumatology	~1M	~\$4B	40-60% MODERATE-TO-SEVERE
SLE Rheumatology	Up to 500K	~\$1B	



Souriors: Evaluate Pharma, Company Estimates, Viral Street research
"Global drug receiver effects to the total market across all sevently levels
Notes: SLE = systemic lupus enythematosus; "Group of indications based on current midfate-stage trials for BMS's allosteric TYK2 inhibitor deucravacitinib; global commercial sales totaled \$10.658 for biologics targeting IL-12/2

Notes: SLE = systemic lupus enythematosus; "Group of indications based on current midfate-stage trials for BMS's allosteric TYK2 inhibitor deucravacitinib; global commercial sales totaled \$10.658 for biologics targeting IL-12/2

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Notes: SLE = systemic IL-12/2

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Notes:



ORALLY BIOAVAILABLE selective allosteric inhibitor of TYK2



VTX958 Program Summary

Allosteric, Selective TYK2 Inhibitor

Potentially Differentiated TYK2 Inhibitor

- · Selective, allosteric TYK2 inhibitor
- TYK2 functional selectivity can potentially differentiate clinical profile vs. less selective TYK2 inhibitors

Clinically Validated Target

- Well established clinical efficacy in psoriasis, IBD and psoriatic arthritis with biologics targeting IL-12/IL-23 and IL-23* pathways
- These pathways also the target of allosteric TYK2 inhibitors
- Phase 3 PoC in psoriasis has been demonstrated** by BMS' allosteric TYK2 inhibitor deucravacitinib

Deucravacitinib in Phase 2/3 for Crohn's disease, psoriatic arthritis, lupus

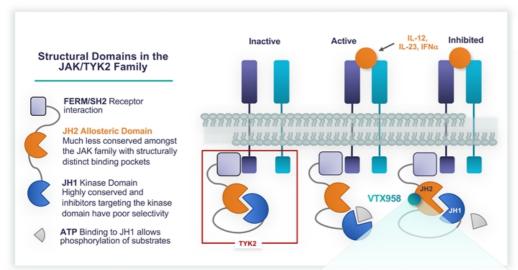
Large Addressable Markets

 Multiple autoimmune disorders in dermatology, IBD, renal and rheumatology total \$45B WW



Includes approved drugs Stelara™ (.NU), Tremfya® (.INU), Skyrizi™(ABBV), Ilumya™ (Sun Pharma) and others in late-stage development (mirikizumab (LLY), brazikumab (AZN)
**Deucravactinib efficacy reported on 16-week primary endopoint of PASI-75 (75% reduction of psoriasis affected area and severity) at AAD 27:
p-0.0001 vs. placebo and Otecalib in POETYK-1; p-0.003 vs. Otecal in POETYK-1; Ses side 14 for more detail on S458 workdwide market

Allosteric Inhibitor VTX958 Binds Selectively to the TYK2 JH2 Domain



Targeting the JH2 (allosteric)
domain of TYK2 affords TYK2
inhibitors with selectivity against other
JAK isoforms

Features of VTX958 JH2 Allosteric Inhibition

Selectivity for TYK2 JH2 vs. JAK1 JH2 domain (>4,000 X)

No binding to JAK2/3 JH2 domains

No binding to TYK2 kinase JH1 and

No kinase enzyme inhibition of any JAK family member

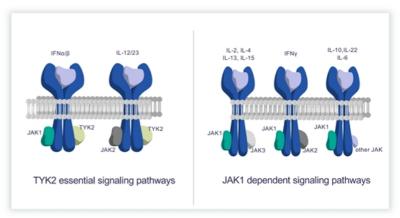
Selective TYK2 Inhibitor



VTX958 More Selective than Deucravacitinib for TYK2 JH2 Domain

Inhibits TYK2 Pathways (IL-12, IL-23, IFNα) while Avoiding the JAK1/2/3 Pathways

	DEUCRAVACITINIB	VTX958
TYK2-JH2 Binding K _d	0.009 nM	0.058 nM
JAK1-JH2 Binding K _d	0.43 nM	240 nM
Selectivity (fold)	48	>4,000





Source: Ventyx internal data

VTX958 Selectively Targets IL-12, IL-23 and IFNα

VTX958 Potently Inhibits TYK2 Pathways

Selective and potent inhibition of IL-12/23 and Type I interferon axis allows targeting pathways driving immunemediated diseases

PROINFLAM	MATORY INNATE 8	тн1/тн17 сүто	KINES
	P	soriasis Patient PBM	ic
DRUG	IL-12 IC ₅₀ (nM)	IL-23 IC ₅₀ (nM)	IFNα IC ₅₀ (nM)
VTX958	35	5	12
deucravacitinib	10	10	5

VTX958 Has No Measurable Inhibition of JAK1-Mediated Pathways

Lack of inhibition of IL-6, IL-10 and other protective cytokines may avoid potential AEs associated with less selective inhibitors

PLEIOTROPIC CYTOKINES WITH PROTECTIVE FUNCTIONS						
DRUG	IL-22 IC ₅₀ (nM)	IL-10 IC ₅₀ (nM)	IFNγ IC ₅₀ (nM)	IL-4 IC ₅₀ (nM)	IL-6 IC ₅₀ (nM)	
VTX958	>10,000	>10,000	>10,000	>10,000	>10,000	
deucravacitinib	114	20	350	249	464	



- · Potent activity against IL-23, a key cytokine implicated in psoriasis and other indications
- Broad therapeutic window with VTX958 may allow for higher exposures in Phase 2/Phase 3 studies



Source: Ventyx internal data; conducted in peripheral blood mononuclear cells (PBM

VTX958 Phase 1 SAD Results Support Clinical Advancement

Safety

Well-tolerated across all cohorts; all AEs observed were mild and not dose- or time-of-dose dependent

Pharmacokinetics

No dose-saturation observed; PK and absorption profiles suggest continued absorption throughout GI tract

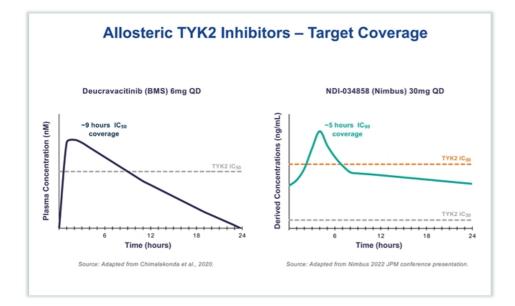
Pharmacodynamics

Dose-dependent VTX958-mediated effect on TYK2 signaling observed in both *in vivo* gene expression studies and *ex vivo* stimulation assays



NOTE: SAD = single ascending dose; AE= adverse event; dose-related exposures are observed at all doses

Targeting a Best-in-Class Exposure Profile With VTX958



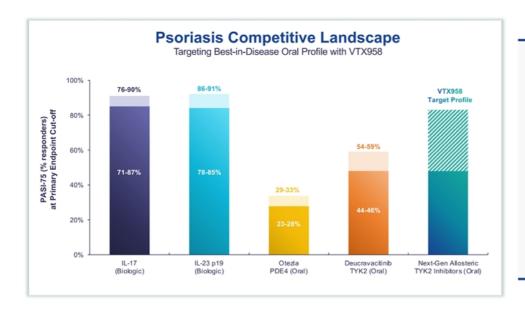
VTX958 Target Profile

- Maximize TYK2 pathway suppression (IC₅₀ and IC₉₀) with once-daily oral dosing
- Wide safety margin enabling higher doses and exposures:
 - Potential for improved efficacy in PsO + PsA with greater TYK2 inhibition
 - Higher exposures may be necessary to achieve efficacy in Crohn's disease



Note: Exposure curves are adapted from corporate presentations and publications, as noted

VTX958 Profile Expected to Drive Clinical Differentiation



KEY TAKEAWAYS

- Current oral options in PsO are substantially less efficacious than biologics
- Greater TYK2 suppression may produce improved efficacy compared to other oral agents, with potential to approach leading biologics
- Significant opportunity for a best-in-disease oral agent in psoriasis, a >\$20B global market



Note: Solid area represents pbo-adjusted response rate; dashed area indicates total observed response rate; primary endpoint cut-off ranges from Week 10 to Week 16 Sources: Company reports and FDA labels for approved anti-IL-17 and anti-IL-23 biologics

Unlocking the Opportunity in Crohn's Disease

Several-fold Higher Doses Required in Crohn's*

Agent	PsO Dose	Crohn's Dose
Skyrizi (IL-23)	150mg Q12W Subcutaneous	600mg IV Q4W (induction) 360mg SC (maintenance)
Tremfya (IL-23)	100mg Q8W Subcutaneous	200mg / 600mg / 1200mg** IV Q4W induction
Stelara (IL-12/23)	40mg / 90mg Q12W Subcutaneous	260mg / 390mg / 520mg IV induction dose
Humira (TNFα)	80mg (SC induction) 40mg Q2W maintenance	160mg (SC induction) 40mg Q2W maintenance

Greater Exposures Needed for TYK2 Inhibitor Efficacy in Crohn's

- Biologics data suggest substantially higher exposures are required for efficacy in Crohn's vs. PsO
- Maximizing TYK2 target coverage expected to differentiate VTX958 from other TYK2 inhibitors
- Selectivity, safety and tolerability considerations may limit the Crohn's opportunity for other TYK2 inhibitors
- Optimized profile of VTX958 may unlock a major market opportunity in Crohn's, a >\$13B global market



Source: FDA labels for approved drugs/indications; Skyrizi represents dose submitted for FDA approval. Note: maintenance dose unless otherwise specified.
**Represents Phase 2 doses; specific Phase 3 doses not disclosed.

VTX958 Clinical Development Plan





NOTE: SAD = single ascending dose: MAD = multiple ascending dose: PoC = proof-of-conce



PERIPHERALLY RESTRICTED S1P1R MODULATOR

with potential for treatment of ulcerative colitis



VTX002 Program Summary

Phase 2 S1P1R Modulator for Ulcerative Colitis

Potentially Differentiated S1P1R Modulator

- Selective S1P1R modulator
- Differentiated on key parameters
- Demonstrated pharmacodynamic activity in Phase 1 trial
- Pursuing clinical development plan in both treatment-naïve and biologic-experienced patients

Clinically Validated Target

- S1P1R modulators approved for MS and UC with clinical trials ongoing in other indications
- BMS' ozanimod approved for UC in May 2021

Large Addressable Markets

 Ulcerative colitis is lead indication totaling up to \$7B in worldwide revenue



VTX002 Differentiates on Multiple Key Parameters vs. Competitors

Potential for Differentiated Clinical Profile in UC Patients

Sustained lymphocyte reduction up to 65% across multiple doses in MAD trial

Safety Profile

No SAEs, elevated LFTs, abnormal PFTs or macular edema

No Drug-Drug Interactions

No CYP inhibition; no food effect; favorable profile for patients with co-morbidities

Fast Onset of Action Faster Lymphocyte Recovery

No long-acting circulating metabolites Optimal half life (t~20h)

Ability to Dose Titrate

Potential to avoid first-dose cardiac monitoring in label

Peripherally Restricted

Very low CNS penetration; not a repurposed MS drug; potential to avoid macular edema



Notes: SAE=significant adverse event; MAD=multiple ascending dose

VTX002 Differentiates on Multiple Key Parameters vs. Competitors

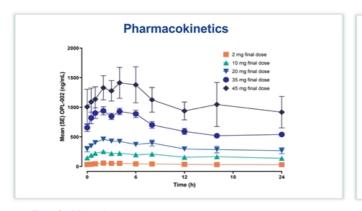
Differentiating Parameter	Ozanimod	Etrasimod	VTX002
Receptor Selectivity	S1P 1,5	S1P 1,4,5	S1P 1,5
Lymphocyte Suppression in Healthy Volunteers	1 mg, ~60%	2 mg, 69%	40 mg, ~65%
Lymphocyte Suppression in UC Patients*	1_mg, 49%	2 mg, 40%	TBD
CYP450 Interactions	Yes	No	No
Liver Enzyme Elevations	Yes	No	No
Active Metabolites	Yes	No	No
Half-life	19 h, Met 10-13 d	33 h	~20 h
Fast Lymphocyte Recovery Time	No	Yes	Yes
First Dose Heart Rate Reduction	Yes	Yes	Yes
Dose Titration	Yes	No	Yes
First Dose Monitoring	No	TBD	TBD



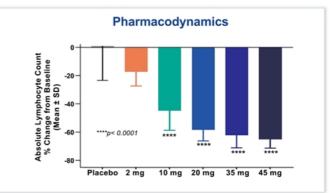
Source: NEJM (2016), Gastroenterology (2020)
*Ph2 UC ALC reduction from baseline: 1mg ozanimod (49%), 2mg etrasimod (40%)

Phase 1 MAD Results: Dose-Dependent Exposure and Lymphocyte Reduction

Absolute Lymphocyte Count (ALC) Reductions of 40-50% Correlated with Clinical Efficacy Observed in UC*



- T_{1/2} of ~20 hours
- Dose-proportionate exposure after single and multiple doses of VTX002 with steady-state reached after 4 to 7 days of targetdose exposure



 Demonstrated consistent, sustained reduction of lymphocytes up to 65% across multiple dose groups



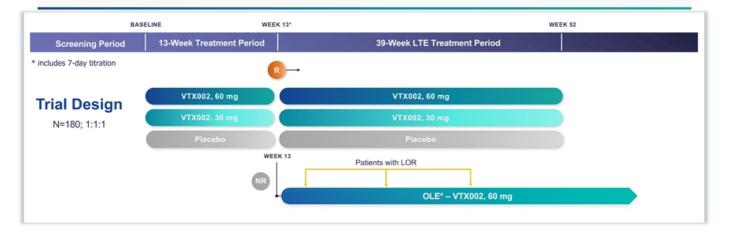
Source: NEJM (2016), Gastroenterology (2020)

*Ph2 UC ALC reduction from baseline: 1mg ozanimod (49%), 2mg etrasimod (40%)

Phase 2 Induction Trial in Moderate-to-Severe Ulcerative Colitis

KEY TAKEAWAYS

- · Powered for primary endpoint of clinical remission
- · Trial may serve as the first of two pivotal trials required for registration

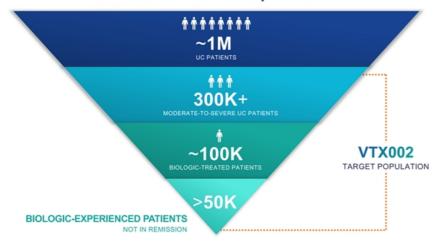




Note: Phase 2 tablet doses of 30mg and 60mg provide comparable VTX002 exposure as Phase 1 suspension doses of 20mg and 40mg, respectively *Induction and OLE non-responder dosing includes 7-day titration period followed by 12 weeks of placebo or VTX002 dose

Underpenetrated Market for Biologic Refractory Patients

Addressable UC Patient Population in US



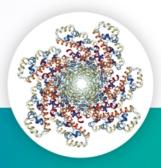
- Existing agents leave room for new treatments
- Novel oral agents may expand penetrance of treated moderate-tosevere UC population beyond current ~25-30%
- S1P well positioned to emerge as leading oral therapeutic class based on its attractive class efficacy/safety profile



2.7



SELECTIVE NLRP3 INFLAMMASOME INHIBITORS for systemic and CNS indications



Rationale for Targeting the NLRP3 Inflammasome

NLRP3 Inflammasome Inhibitors Target IL-1β, a Key Driver of Inflammatory Disease

In vivo Evidence

- The NLRP3 inflammasome can become overactive in the presence of persistent tissue damage or crystal deposits
- Inflammasome activation results in release of IL-1β & IL-18 recruiting neutrophils and driving Th17 response
- This leads to pyroptosis and further tissue damage

Genetic Evidence

 Gain-of-function mutations in the NLRP3 gene, associated with certain severe orphan inflammatory diseases, are classified as cyropyrin-associated periodic syndromes (CAPS)

Clinical Validation of Downstream Target

- IL-1β signaling, downstream of inflammasome activation, is a clinicallyvalidated, anti-inflammatory target with biologics
- Ilaris® (\$873M sales in 2020) approved for CAPS and other orphan periodic fever syndromes



NLRP3 = NOD-like receptor family, pyrin domain-containing protein 3; IL-1β = interleukin-1β

NLRP3 Inhibitor Program Summary

Peripheral NLRP3 Inhibitor: VTX2735

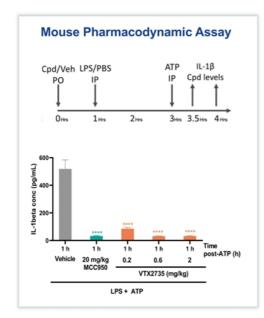
- · Selective NLRP3 inhibitor
- · Well tolerated in GLP safety and tox assessment
- Phase 1 ongoing, expected to complete dosing in H1 2022
- High oral bioavailability in non-clinical PK studies
- PD activity demonstrated in animal models

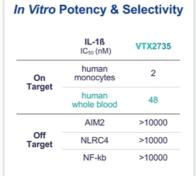
CNS NLRP3 Inhibitor: VTX3232

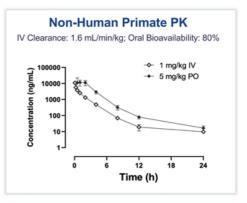
- Selective compounds generated with high CNS bioavailability
- Novel and proprietary lead series
- Plan to submit IND fourth Q4 2022
- Potential to be first, truly CNS-directed NLRP3 inhibitor to enter clinic



VTX2735 is a Selective & Orally Bioavailable NLRP3 Inhibitor







KEY TAKEAWAYS

- · Well-tolerated preclinically in IND-enabling GLP studies
- Oral bioavailability (80%) in NHP and dose-proportional exposure that predicts potential for wide safety margins based on PK/PD modeling



MCC950 is an NLRP3 inhibitor and a control compound used in in vitro and in vivo studie

VTX2735 Has Broad Activity Against Multiple NLRP3 Mutations

Potential for Differentiation in CAPS Setting*

What is CAPS?

An ultra-orphan auto-inflammatory disease caused by various mutations in NLRP3 and characterized by inappropriate release of IL-1β and symptoms of recurrent systemic inflammation

KEY TAKEAWAY

VTX2735 blood assay data from CAPS patients suggest inhibitory activity across several mutations: FCAS, MWS and NOMID subset of CAPS patients

			75% II CAPS pati North Amer				MOST SEVERE
CPD	CHALLENGE	FCAS1 L353P	FCAS2 (L353P)	FCAS3 (L353P)	FCAS4 A439V/G564R	FCAS.MWS E525K/V198M	NOMID F309Y
VTX2735	LPS	117	56	166	14	24	17
MCC950	LPS	>10K	>10K	>10K	1,264	>10K	>10K

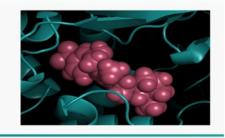


*Source: UCSD (Dr. Hal Hoffman's lab); CAPS=Cryopyrin-Associated Periodic Syndrome

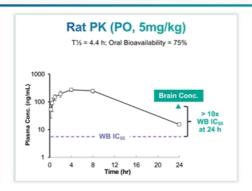
CNS-Penetrant NLRP3 Inhibitor VTX3232

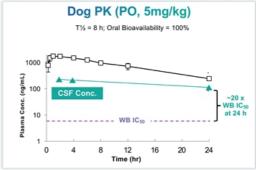
KEY TAKEAWAYS

- · Novel, potent, brain-penetrant inhibitor of NLRP3
- 13 nM IC₅₀ in human whole blood IL-1β release assay
- · Unique structural chemotype vs. peripheral NLRP3 inhibitors
- Provisional application filed June 2021
- IND filing in Q4 2022; Phase 1 in Q1 2023



	In Vitro Potency & Selectivity				
	IL-1β VTX323 Assay IC ₅₀ (nM				
NLRP3	huWB	13			
AIM2		>10000			
NLRC4	вмрм	>10000			
NF-kb		>10000			







NLRP3 Program Clinical Development Plan







Our Comprehensive NLRP3 Portfolio Targets a Broad Range of Major Inflammatory Diseases

NLRP3

Systemic Diseases

Peripheral NLRP3 inhibitors are designed to treat cardiovascular, rheumatic, fibrotic and rare genetic diseases





- Cardiovascular
- Rheumatic
- Fibrotic Diseases
- · Rare Genetic Diseases

Our solution: VTX2735

Neuroinflammatory Diseases

CNS-directed NLRP3 inhibitors are designed to treat a range of neurodegenerative disorders, such as Alzheimer's and Parkinson's disease



- · Alzheimer's Disease
- · Parkinson's Disease
- ALS

Our solution: VTX3232





PROJECTED PIPELINE CATALYSTS AND SUMMARY

3.3

Projected Catalysts Over Next 24 Months

	PROGRAMS		PROGRAMS		H1'2021 H2'2021 H1'2022		H2'2022	2023
Sept.	VTX958 Allosteric TYK2 inhibitor addressing a broad range of autoimmune disorders	Phase	1 SAD	Phase 1 MAD	Phase 2 in	Multiple Indications*		
	VTX002 Selective S1P1R modulator targeting UC and other immune disorders			Phase 2 Ulcerative	Colitis 13-Week I	nduction		
STATE OF THE PARTY	VTX2735 Peripheral NLRP3 inflammasome inhibitor for multiple inflammatory and immune conditions	IND-ena	bling	Phase 1 SAD/MAD	Phase 2	PoC Initiation		
	VTX CNS CNS-directed NLRP3 inflammasome inhibitor for neurodegenerative diseases	Candida	ate Selection	IND-ena	bling	Phase 1 SAD/MAD**		



^{*}Following completion of our Phase 1 trial, we intend to initiate Phase 2 PoC trials in psoriasis, psoriatic arthritis, Crohn's disease and potentially other indications
** Following regulatory acceptance of planned Q4 2022 IND filing, we intend to initiate and conduct a Phase 1 SAD/MAD trial in healthy volunteers

Investment Highlights

PRODUCTIVE IMMUNOLOGY PLATFORM

- · Internal R&D engine designed to generate candidates to address autoimmune and inflammatory diseases with high unmet need
- 100% commercial rights to entire portfolio; long patent life for all product candidates

POTENTIALLY DIFFERENTIATED MEDICINES

- Multiple selective, oral, small molecule product candidate portfolio:
 - VTX958: allosteric TYK2 inhibitor for multiple autoimmune indications
 - VTX002: peripherally-restricted S1P1R modulator for ulcerative colitis
 - VTX2735: peripheral NLRP3 inhibitor for multiple autoimmune indications
 - VTX3232: CNS-targeted NLRP3 inhibitor for multiple neurodegenerative indications

TARGET MAJOR INFLAMMATORY & IMMUNOLOGY DISEASE MARKETS

- · Our portfolio can address I&I markets, such as psoriasis, IBD, and other indications
- · Opportunity to disrupt existing markets dominated by biologics with varying degrees of efficacy and safety in order to:
 - Capture refractory patients
 - Expand market share of moderate-to-severe patient populations with patient-friendly oral therapy

STRONG BALANCE SHEET

- · Over \$339 million raised from IPO and dedicated biotech investors
- · Cash balance of \$142M as of September 30, 2021*



*Not including gross proceeds of \$174M raised in October 2021 IPO

