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 6, 2023

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 8.01 Other Events.

On March 6, 2023, Aclaris Therapeutics, Inc. issued a press release announcing the preliminary topline data for its Phase 2a clinical trial of zunsemetinib, an investigational oral MK2 inhibitor, in subjects with moderate to severe hidradenitis suppurativa and related matters. A copy of this press release is filed as Exhibit 99.1 to this Current Report on Form 8-K and is incorporated herein by reference.

Item 9.01 Financial Statements and Exhibits.

(d) Exhibits

Exhibit Number	Exhibit Description
99.1	Press Release dated March 6, 2023.
104	The cover page from Aclaris Therapeutics, Inc.'s Form 8-K filed on March 6, 2023, formatted in Inline XBRL.

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/ Douglas Manion
Douglas Manion
Chief Executive Officer and President

Date: March 6, 2023

Aclaris Therapeutics Announces Preliminary Topline Data from 12-Week Phase 2a Study of Oral Zunsemetinib (ATI-450) for Moderate to Severe Hidradenitis Suppurativa

- Study Did Not Meet Primary or Secondary Efficacy Endpoints in Hidradenitis Suppurativa
- Overall Safety Profile and PK/PD Generally Consistent with Observations in Prior Studies of Zunsemetinib

WAYNE, Pa., Mar. 6, 2023 (GLOBE NEWSWIRE) – Aclaris Therapeutics, Inc. (NASDAQ: ACRS), a clinical-stage biopharmaceutical company focused on developing novel drug candidates for immuno-inflammatory diseases, today announced preliminary topline results from a 12-week, Phase 2a, multicenter, randomized, placebo-controlled clinical study to investigate the efficacy, safety, tolerability, pharmacokinetics (PK) and pharmacodynamics (PD) of zunsemetinib (ATI-450), an investigational oral MK2 inhibitor, in patients with moderate to severe hidradenitis suppurativa (HS) (ATI-450-HS-201).

Efficacy
The study did not meet its primary endpoint of change from baseline in inflammatory nodule/abscess count (AN) of zunsemetinib 50mg BID versus placebo at week 12. The study also did not meet the secondary efficacy endpoints assessed in the topline data, including percentage of patients achieving HiSCR-50. The placebo effect observed across all efficacy endpoints was higher than what has been observed in other published HS studies reported to date.

Safety
Zunsemetinib was generally well tolerated. Safety findings were generally consistent with observations from prior clinical studies of zunsemetinib. The most common treatment-emergent adverse events in patients treated with zunsemetinib were dizziness (16.7%), diarrhea (12.5%), headache (12.5%), creatine phosphokinase (CPK) elevation (10.4%) and acne (10.4%), with a majority deemed mild to moderate in severity. Dizziness, diarrhea, headache and CPK elevation were generally transient in nature. Thirty-seven patients discontinued study treatment (22 on zunsemetinib and 15 on placebo), with 15 patients discontinuing treatment due to AEs (11 on zunsemetinib and 4 on placebo). No serious adverse events and no serious and/or opportunistic infections were observed with patients treated with zunsemetinib.

Pharmacokinetics/Pharmacodynamics
PK and PD were generally consistent with observations from prior clinical studies of zunsemetinib. A preliminary analysis of endogenous plasma cytokines and chemokines in patients with a confirmed dose of study treatment on the day of blood draw, demonstrated zunsemetinib dependent inhibition relative to placebo. Of the proinflammatory markers that were elevated at baseline relative to healthy donors, including IL6, IL8 and MIP1b, treatment-related median inhibition trends were observed across the 12-week dosing period. While the proinflammatory cytokine IL12/23p40 was not elevated at baseline relative to healthy donors, a median inhibition trend was observed in patients treated with zunsemetinib. In the subset of patients with quantifiable IL17A/F levels at baseline, the cytokine was elevated relative to healthy donors and an inhibition trend was observed with zunsemetinib treatment. Endogenous TNFα plasma levels were not elevated relative to healthy donors yet a small inhibition trend was observed in treated patients. The anti-inflammatory cytokine IL1RA was elevated at baseline and treatment-related inhibition was not observed.

“We are incredibly grateful to the patients, investigators, our internal study team and all others who contributed to the execution of this clinical trial,” stated Doug Manion, M.D. Aclaris’ Chief Executive Officer. “Despite not producing the efficacy results we had hoped for in this particularly challenging disease, we are encouraged by the consistent demonstration of zunsemetinib’s mechanism of action and the strengthening of our safety data base and continue to look forward to our next data read out of our Phase 2b study of zunsemetinib in patients with moderate to severe rheumatoid arthritis.”

About ATI-450-HS-201 Phase 2a Study (NCT05216224)

ATI-450-HS-201 is a randomized, double-blind, placebo-controlled, multi-center Phase 2a study to investigate the efficacy, safety, tolerability, PK and PD of zunsemetinib (50 mg twice daily) in patients with moderate to severe HS. The study enrolled 95 patients who were randomized 1:1 to zunsemetinib (n=48) or placebo (n=47) across 19 clinical study sites in the US. The primary endpoint of the study is change from baseline in inflammatory nodule/abscess count at week 12. The topline efficacy analyses are based on the full analysis set. Healthy volunteer samples for PD analyses were obtained outside of the clinical trial setting.

About Zunsemetinib (ATI-450)

Zunsemetinib is an investigational oral mitogen-activated protein kinase-activated protein kinase 2 (MK2) inhibitor. This mechanism potentially leads to the inhibition of multiple cytokines, chemokines, matrix metalloproteases and other inflammatory signals. Key inflammatory cytokines driven by this mechanism include TNFα and interleukin-1α, -1β, -6, -8 and -17 (IL1α, IL1β, IL6, IL8 and IL17). Aclaris is developing zunsemetinib as a potential treatment for rheumatoid arthritis and psoriatic arthritis.

About Aclaris Therapeutics, Inc.

Aclaris Therapeutics, Inc. is a clinical-stage biopharmaceutical company developing a pipeline of novel drug candidates to address the needs of patients with immuno-inflammatory diseases who lack satisfactory treatment options. The company has a multi-stage portfolio of drug candidates powered by a robust R&D engine exploring protein kinase regulation. For additional information, please visit www.aclaristx.com.

Cautionary Note Regarding Forward-Looking Statements

Any statements contained in this press release 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,” “intend,” “may,” “plan,” “potential,” “will,” and similar expressions, and are based on Aclaris’ current beliefs and expectations. These forward-looking statements include Aclaris’ expectations regarding the timing of reporting results from its clinical studies. 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, Aclaris’ ability to enter into strategic partnerships on commercially reasonable terms, the uncertainty regarding the macroeconomic environment and the COVID-19 pandemic and other risks and uncertainties that are described in the Risk Factors section of Aclaris’ Annual Report on Form 10-K for the year ended December 31, 2022 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” page of the “Investors” section of Aclaris’ website at www.aclaristx.com. Any forward-looking statements speak only as of the date of this press release and are based on information available to Aclaris as of the date of this release,

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.

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