

**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): September 26, 2023

**TALARIS THERAPEUTICS, INC.**  
(Exact name of registrant as specified in its charter)

**Delaware**  
(State or other jurisdiction  
of incorporation)

**001-40384**  
(Commission  
File Number)

**83-2377352**  
(I.R.S. Employer  
Identification No.)

**93 Worcester St.**  
**Wellesley, Massachusetts**  
(Address of principal executive offices)

**02481**  
(Zip Code)

Registrant's telephone number, including area code: (502) 398-9250

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	Trade Symbol(s)	Name of each exchange on which registered
Series A Common Stock, \$0.0001 par value per share	TALS	The Nasdaq Global 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

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

As previously announced, on June 22, 2023, Talaris Therapeutics, Inc., a Delaware corporation (“Talaris”), Terrain Merger Sub, Inc., a Delaware corporation and a wholly owned subsidiary of Talaris (“Merger Sub”), and Tourmaline Bio, Inc., a Delaware corporation (“Tourmaline”), entered into an Agreement and Plan of Merger (the “Merger Agreement”), pursuant to which, among other matters, and subject to the satisfaction or waiver of the conditions set forth in the Merger Agreement, Merger Sub will merge with and into Tourmaline, with Tourmaline continuing as a wholly owned subsidiary of Talaris and the surviving corporation of the merger (the “Merger”).

On September 26, 2023, Tourmaline updated information reflected in an investor presentation, which is attached as Exhibit 99.1 to this Current Report on Form 8-K and is incorporated herein by reference. Representatives of Tourmaline will use the updated presentation in various meetings with investors and analysts from time to time.

The information in Item 7.01 of this Current Report on Form 8-K, including the information set forth in Exhibit 99.1, is being furnished and shall not be deemed “filed” for purposes of Section 18 of the Securities Exchange Act of 1934, as amended (the “Exchange Act”), nor shall Exhibit 99.1 furnished herewith be deemed incorporated by reference in any filing under the Securities Act of 1933, as amended, or the Exchange Act, except as shall be expressly set forth by specific reference in such a filing.

### Forward-Looking Statements

This Current Report on Form 8-K and the exhibit furnished herewith contain “forward-looking statements” within the meaning of the “safe harbor” provisions of the Private Securities Litigation Reform Act of 1995, including but not limited to, express or implied statements regarding the structure, timing and completion of the proposed Merger; the combined company’s listing on Nasdaq after closing of the proposed Merger; expectations regarding the ownership structure of the combined company; the anticipated timing of the closing of the Merger; each company’s and the combined company’s expected cash position at the closing of the proposed Merger and cash runway of the combined company; the future operations of the combined company; the nature, strategy and focus of the combined company; the development and commercial potential and potential benefits of any product candidates or platform technologies of the combined company; anticipated preclinical and clinical drug development activities and related timelines, including the expected timing for data and other clinical results; the competitive landscape of the combined company; anticipated intellectual property timelines; and other statements that are not historical fact. All statements other than statements of historical fact contained in this Current Report on Form 8-K are forward-looking statements. These forward-looking statements are made as of the date they were first issued, and were based on the then-current expectations, estimates, forecasts, and projections, as well as the beliefs and assumptions of management. There can be no assurance that future developments affecting Talaris, Tourmaline or the proposed transaction will be those that have been anticipated.

Forward-looking statements are subject to a number of risks and uncertainties, many of which involve factors or circumstances that are beyond Talaris’ control. Talaris’ actual results could differ materially from those stated or implied in forward-looking statements due to a number of factors, including but not limited to (i) the risk that the conditions to the closing of the proposed Merger are not satisfied, including the failure to timely obtain shareholder approval for the transaction, if at all; (ii) uncertainties as to the timing of the consummation of the proposed Merger and the ability of each of Talaris and Tourmaline to consummate the proposed Merger; (iii) risks related to Talaris’ ability to manage its operating expenses and its expenses associated with the proposed Merger pending closing; (iv) risks related to the failure or delay in obtaining required approvals from any governmental or quasi-governmental entity necessary to consummate the proposed Merger; (v) the risk that as a result of adjustments to the exchange ratio, Talaris shareholders and Tourmaline stockholders could own more or less of the combined company than is currently anticipated; (vi) risks related to the market price of Talaris’ common stock relative to the value suggested by the exchange ratio; (vii) unexpected costs, charges or expenses resulting from the transaction; (viii) potential adverse reactions or changes to business relationships resulting from the announcement or completion of the proposed Merger; (ix) the uncertainties associated with Tourmaline’s platform technologies, as well as risks associated with the clinical development and regulatory approval of product candidates, including potential delays in the commencement, enrollment and completion of clinical trials; (x) risks related to the inability of the combined company to obtain sufficient additional capital to continue to advance these product candidates and its preclinical programs; (xi) uncertainties in obtaining successful clinical results for product candidates and unexpected costs that may result therefrom; (xii) risks related to the failure to realize any value from product candidates and preclinical

programs being developed and anticipated to be developed in light of inherent risks and difficulties involved in successfully bringing product candidates to market; (xiii) risks associated with the possible failure to realize certain anticipated benefits of the proposed Merger, including with respect to future financial and operating results; (xiv) risks associated with Talaris' financial close process; (xv) the risk that the pre-closing financing is not consummated; and (xvi) the risk that Talaris shareholders receive more or less of the cash dividend than is currently anticipated, among others. Actual results and the timing of events could differ materially from those anticipated in such forward-looking statements as a result of these risks and uncertainties. These and other risks and uncertainties are more fully described in periodic filings with the SEC, including the factors described in the section titled "Risk Factors" in Talaris' Annual Report on Form 10-K for the year ended December 31, 2022 filed with the SEC, and in other filings that Talaris makes and will make with the SEC in connection with the proposed Merger, including the Proxy Statement described below under "Additional Information and Where to Find It." You should not place undue reliance on these forward-looking statements, which are made only as of the date hereof or as of the dates indicated in the forward-looking statements. Talaris expressly disclaims any obligation or undertaking to release publicly any updates or revisions to any forward-looking statements contained herein to reflect any change in its expectations with regard thereto or any change in events, conditions or circumstances on which any such statements are based. This Current Report on Form 8-K does not purport to summarize all of the conditions, risks and other attributes of an investment in Talaris or Tourmaline.

### **Participants in the Solicitation**

This Current Report on Form 8-K and the exhibit filed or furnished herewith relate to the proposed merger transaction involving Talaris and Tourmaline and may be deemed to be solicitation material in respect of the proposed merger transaction. In connection with the proposed merger transaction, Talaris has filed relevant materials with the SEC, including a registration statement on Form S-4 (the "Form S-4") that contains a proxy statement (the "Proxy Statement") and prospectus. This Current Report on Form 8-K is not a substitute for the Form S-4, the Proxy Statement or for any other document that Talaris may file with the SEC and or send to Talaris' shareholders in connection with the proposed merger transaction. **BEFORE MAKING ANY VOTING DECISION, INVESTORS AND SECURITY HOLDERS OF TALARIS ARE URGED TO READ THE FORM S-4, THE PROXY STATEMENT AND OTHER DOCUMENTS FILED WITH THE SEC CAREFULLY AND IN THEIR ENTIRETY WHEN THEY BECOME AVAILABLE BECAUSE THEY WILL CONTAIN IMPORTANT INFORMATION ABOUT TALARIS, THE PROPOSED MERGER TRANSACTION AND RELATED MATTERS.**

### **No Offer or Solicitation**

This Current Report on Form 8-K and the exhibit furnished herewith do not constitute an offer to sell or the solicitation of an offer to buy any securities nor a solicitation of any vote or approval with respect to the proposed transaction or otherwise. No offering of securities shall be made except by means of a prospectus meeting the requirements of Section 10 of the U.S. Securities Act of 1933, as amended, and otherwise in accordance with applicable law.

### **Additional Information and Where to Find It**

Investors and security holders may obtain free copies of the Form S-4, the Proxy Statement and other documents filed by Talaris with the SEC through the website maintained by the SEC at <http://www.sec.gov>. Copies of the documents filed by Talaris with the SEC are also available free of charge on Talaris' website at [www.talaristx.com](http://www.talaristx.com), or by contacting Talaris' Investor Relations at [investors@talaristx.com](mailto:investors@talaristx.com). Talaris, Tourmaline, and their respective directors and certain of their executive officers may be considered participants in the solicitation of proxies from Talaris' shareholders with respect to the proposed merger transaction under the rules of the SEC. Information about the directors and executive officers of Talaris is set forth in its Annual Report on Form 10-K for the year ended December 31, 2022, which was filed with the SEC on March 31, 2023, and in subsequent documents filed with the SEC. Additional information regarding the persons who may be deemed participants in the proxy solicitations and a description of their direct and indirect interests, by security holdings or otherwise, are also included in the Form S-4, the Proxy Statement and other relevant materials to be filed with the SEC when they become available. You may obtain free copies of this document as described above.

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**Item 9.01 Financial Statements and Exhibits**

**(d) Exhibits**

Exhibit No.	Document
99.1	Investor Presentation of Tourmaline Bio, Inc., dated September 26, 2023.
104	Cover Page Interactive Data File (embedded within the Inline XBRL document).



**SIGNATURE**

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.

TALARIS THERAPEUTICS, INC.

Date: September 26, 2023

By: /s/ Mary Kay Fenton

May Kay Fenton

Chief Financial Officer and Interim Chief Financial Officer

# TOURMALINE

**Sell-side Analyst Day**

September 26, 2023

# Disclaimer

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Tourmaline obtained the industry, market and competitive position data used throughout this presentation from its own internal estimates and research, as well as from industry and general publications, and research, surveys and studies conducted by third parties. Internal estimates are derived from publicly available information released by industry analysts and third-party sources, Tourmaline's internal research and its industry experience, and are based on assumptions made by Tourmaline based on such data and its knowledge of the industry and market, which it believes to be reasonable. In addition, while Tourmaline believes the industry, market and competitive position data included in this presentation is reliable and based on reasonable assumptions, Tourmaline has not independently verified any third-party information, and all such data involve risks and uncertainties and are subject to change based on various factors.

This presentation contains trademarks, service marks, trade names and copyrights of Tourmaline and other companies, which are the property of their respective owners. The use or display of third parties' trademarks, service marks, trade name or products in this presentation is not intended to, and does not imply, a relationship with Tourmaline, or an endorsement of sponsorship by Tourmaline. Solely for convenience, the trademarks, service marks and trade names referred to in this presentation may appear with the ©, TM or SM symbols, but such references are not intended to indicate, in any way, that the company will not assert, to the fullest extent under applicable law, their rights or the right of the applicable licensor to these trademarks, service marks and trade name.

# Disclaimer (continued)

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# Agenda

2:00 – 2:05 pm	Introducing our team
2:05 – 2:20 pm	IL-6 renaissance
2:20 – 3:00 pm	TED
3:00 – 3:40 pm	ASCVD
3:40 – 3:50 pm	Key business items
3:50 – 4:30 pm	Q&A and discussion

## Experienced leadership team



**Sandeep Kulkarni, MD**  
Co-founder and  
Chief Executive Officer



**Yung Chyung, MD**  
Chief Medical  
Officer



**Brad Middlekauff, JD**  
Chief Business Officer and  
General Counsel



**Susan Dana Jones, PhD**  
Chief Technology  
Officer



**Kevin Johnson, PhD**  
Chief Regulatory  
Officer



**Ryan Iarrobino**  
Senior Vice President,  
Product Development



**Gerhard Hagn**  
Senior Vice President,  
Head of Commercial & BD



**Dora Rau**  
Senior Vice President,  
Head of Quality

## Key highlights



An IL-6 renaissance is underway: new insights emerging about a broad range of indications where IL-6 may be clinically validated

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TOUR006 offers potential for low volume, infrequent subcutaneous administration

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We are rapidly advancing TOUR006 into mid/late-stage development

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Our team has extensive experience developing and commercializing antibodies for orphan and autoimmune diseases


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Cash runway expected to fund development through 2026\*

## Our lead indications

### Thyroid eye disease (TED): an inflammatory disease that affects the tissue surrounding the eye

- TOUR006's upstream mechanism of action coupled with its convenient low volume, low frequency, subcutaneous administration profile could make it an optimal treatment option for first-line TED
- Mechanism clinically validated after >300 TED patients treated with IL-6 blockers, showing autoantibody reductions and evidence of clinical benefit
-  Phase 2b TED study expected to begin in Q3 2023

### Atherosclerotic cardiovascular disease (ASCVD): a leading cause of global morbidity and mortality

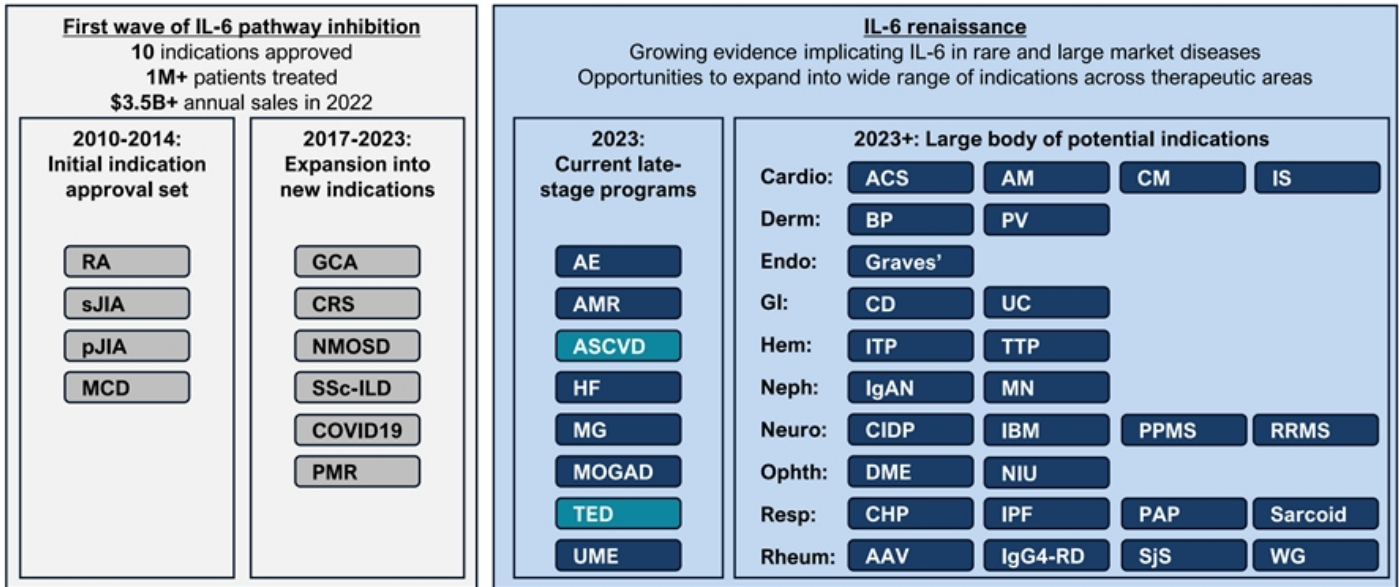
- Emerging clinical evidence appears to validate decades-long research on IL-6 as a key cardiovascular risk factor
- TOUR006 could pursue a fast follower strategy, with potential for less frequent dosing than competitor IL-6 agents in ASCVD
- Phase 2 ASCVD biomarker trial expected to begin in 2024
- External pipeline of phase 3 trials by big pharma has potential to validate IL-6 inhibition in addressing ASCVD and other cardiac disorders





# We are in an IL-6 renaissance

Development timeline for IL-6 pathway inhibitors



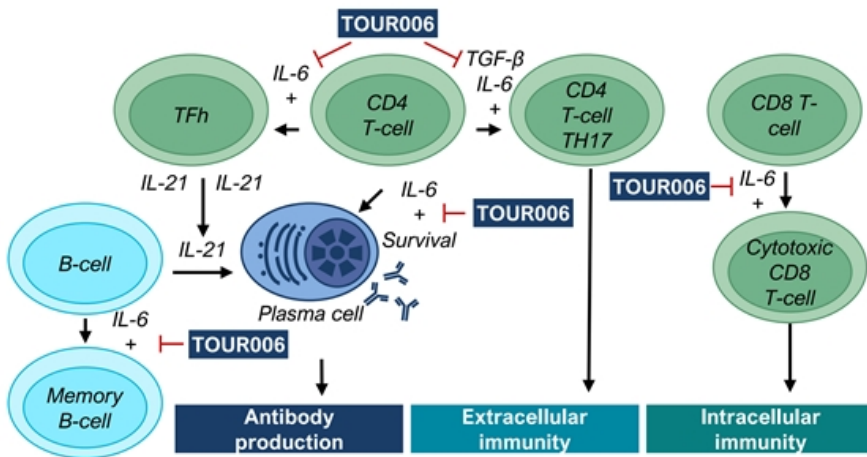
■ Tourmaline indication

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AAV: ANCA-associated vasculitis; ACS: Acute coronary syndrome; AE: Autoimmune encephalitis; AM: Acute myocarditis; AMR: Antibody mediated rejection; ASCVD: Atherosclerotic disease; BP: Bullous pemphigoid; CD: Crohn's disease; CHP: Chronic hypersensitivity pneumonitis; CIDP: Chronic inflammatory demyelinating polyneuropathy; CM: Cardiomyopathy; COVID19: Coronavirus disease 2019; CRS: Cytokine release syndrome; DME: Diabetic macular edema; GCA: Giant cell arteritis; GD: Graves' disease; HF: Heart failure; IBM: Inclusion body myositis; IgAN: IgA nephropathy; IgG4-RD: IgG4 related disease; IPF: Idiopathic pulmonary fibrosis; IS: Ischemic stroke; ITP: Idiopathic thrombocytopenic purpura; MCD: Multicentric castlemans disease; MG: Myasthenia gravis; MN: Membranous nephropathy; MOGAD: Myelin oligodendrocyte glycoprotein antibody-associated disease; NIU: Non-infectious uveitis; NMOSD: Neuromyelitis optica spectrum disorder; PAP: Pulmonary alveolar proteinosis; pJIA: Polyarticular juvenile idiopathic arthritis; PMR: Polymyalgia rheumatica; PPMS: Primary progressive multiple sclerosis; PV: Pemphigus vulgaris; RA: Rheumatoid arthritis; RRMS: Relapsing remitting multiple sclerosis; Sarcoid: Sarcoidosis; sJIA: Systemic juvenile idiopathic arthritis; SJS: Sjögren's syndrome; SSc-ILD: Systemic sclerosis interstitial lung disease; TED: Thyroid eye disease; TTP: Thrombotic thrombocytopenic purpura; UC: Ulcerative colitis; UME: Uveitic macular edema; WG: Wegener's granulomatosis

# IL-6 drives production of autoantibodies and inflammation

## IL-6 mediated impacts on B and T cell pathways<sup>1</sup>



## Translational evidence

- IL-6 enhances antibody production and induces plasma cell differentiation and survival<sup>2</sup>
- In *ex vivo* experiments using samples from patients with NMO, IL-6 shown to promote plasmablast survival and stimulate anti-AQP4 secretion<sup>3</sup>
- Extensive observations in TED and other autoantibody disease that IL-6 blockade suppresses autoantibody levels
- Recent approval of satralizumab in NMOSD offers strong evidence of anti-IL-6's potential in autoantibody driven diseases

# TOUR006, a fully human, high affinity antibody that neutralizes IL-6 is in advanced stages of development

## Fully human antibody that neutralizes IL-6 levels with high affinity

- Kd of 6 pM
- Terminal half-life 47-58 days
- Generated from Medarex transgenic mouse platform

## Robust existing clinical data package

- Two Phase 2 studies completed (SLE and Crohn's)
- 448 subjects have been dosed with TOUR006

## Durable and deep IL-6 signaling blockade observed with infrequent dosing as low as 10mg every 4 weeks

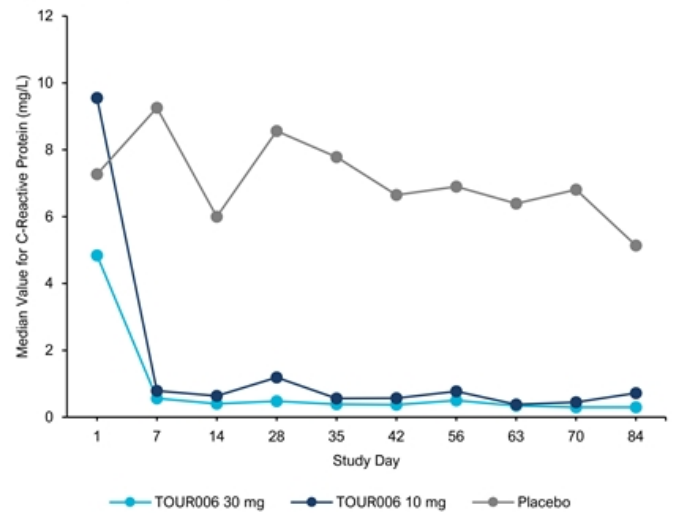
- As measured by C-reactive protein (CRP), a pharmacodynamic marker of IL-6 signaling

## Limited immunogenicity

- Across 448 subjects dosed with TOUR006, only 2 subjects generated ADAs following treatment

## Generally well-tolerated profile to date consistent with IL-6 class

Median serum concentration time profile of CRP from all subjects following day 1, 28, and 56 following multiple intravenous doses of TOUR006 to RA subjects (Study B0151002)



# TOUR006's potential profile: subcutaneous, low volume, low frequency injections

Drug Properties							Dosing interval (days)				
ROA	Drug	Company	Target	Stage	Indications being pursued	Black box warning	QW	Q2W	Q4W	Q8W	Q12W
SC	TOUR006	TOURMALINE	IL-6	In Phase 2b	TED, ASCVD	Drug not approved	56-84				
	Actemra (tocilizumab)	Roche	IL-6R	Approved	RA, GCA, pJIA, sJIA, SSc-ILD	Yes	7-14				
	Kevzara (sarilumab)	REGENERON	IL-6R	Approved	RA, PMR	Yes	14				
	Enspryng (satralizumab)	Roche	IL-6R	Approved	NMOSD, AE, MG, MOGAD, TED	No	28				
	Ziltivekimab	novo nordisk	IL-6	In Phase 3	ASCVD (CKD), HF	Drug not approved	28				
	Clazakizumab	CSL	IL-6	In Phase 2/3	AMR	Drug not approved	28				
IV	Actemra (tocilizumab)	Roche	IL-6R	Approved	RA, GCA, pJIA, sJIA, CRS, COVID19	Yes	28				
	Sylvant (siltuximab)	EUSA Pharma	IL-6	Approved	MCD	No	21				
	Clazakizumab	CSL	IL-6	In Phase 2/3	ASCVD (ESKD)	Drug not approved	28				
IVT	RG6179	Roche	IL-6	In Phase 3	UME, DME	Drug not approved	28				

Indication Key: Approved Investigational

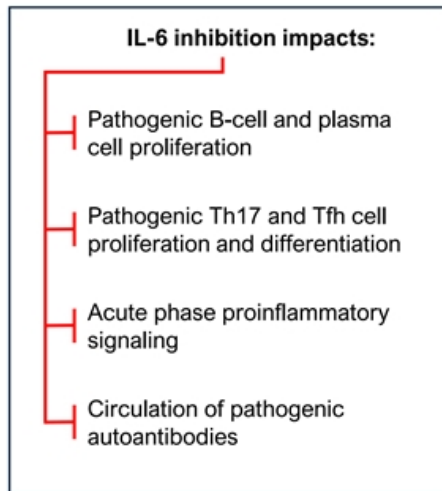
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Source: company reports, publications, FDA review documents, package inserts  
 AE: Autoimmune encephalitis; AMR: Antibody mediated rejection; ASCVD: Atherosclerotic disease; COVID19: Coronavirus disease 2019; CRS: Cytokine release syndrome; DME: Diabetic macular edema; GCA: Giant cell arteritis; HF: Heart failure; MCD: Multicentric castelman's disease; MG: Myasthenia gravis; MOGAD: Myelin oligodendrocyte glycoprotein antibody-associated disease; NMOSD: Neuromyelitis optica spectrum disorder; pJIA: Polyarticular juvenile idiopathic arthritis; PMR: Polymyalgia rheumatica; RA: Rheumatoid arthritis; sJIA: Systemic juvenile idiopathic arthritis; SSc-ILD: Systemic sclerosis interstitial lung disease; TED: Thyroid eye disease; UME: Uveitic macular edema

# TOUR006 has broad potential beyond autoantibody reduction

## An “FcRn-plus” opportunity

### Modes of action for IL-6 inhibition<sup>1,2</sup>

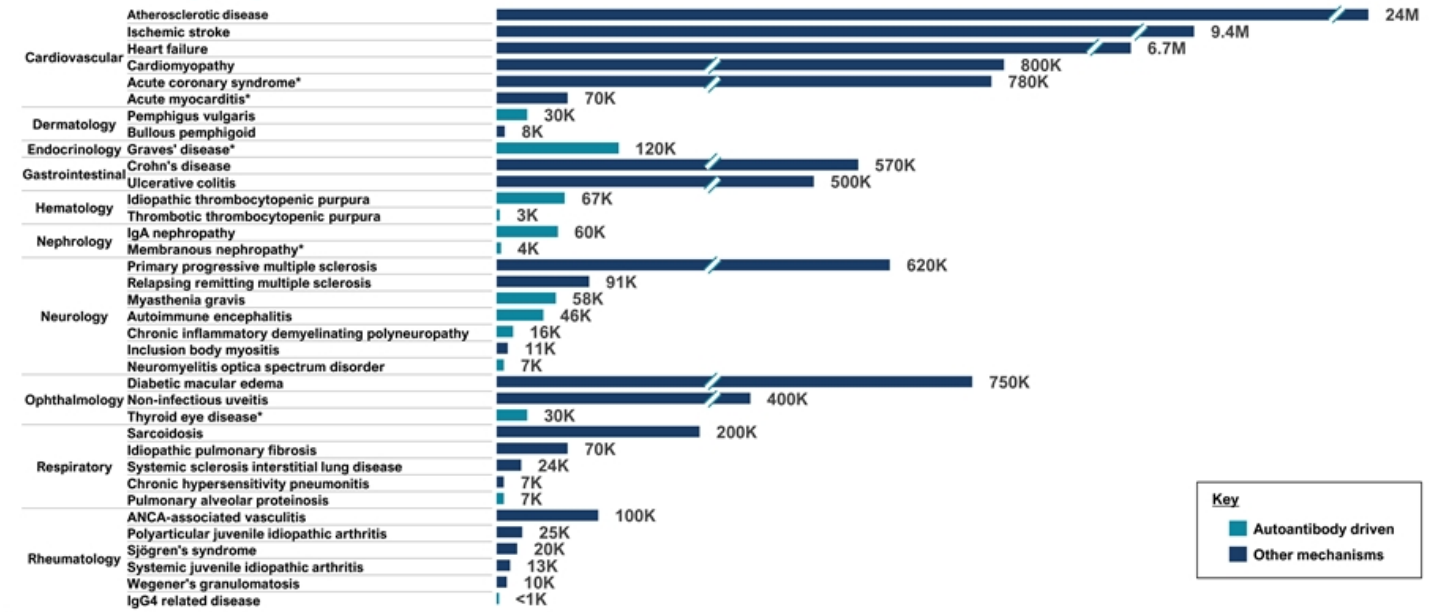


### Potential benefits of IL-6 inhibition versus FcRn inhibition

	FcRn inhibition <sup>3,4,5</sup>	IL-6 inhibition <sup>1,2,6</sup>
Autoantibody reductions	✓	✓
Inhibition of autoantibody production	✗	✓
Anti-inflammatory effects beyond autoantibody reduction	✗	✓
Durability of effect	✗	✓
Low administration burden	✗	✓
Favorable long-term safety profile	?	✓

# IL-6 inhibition has the potential to address a wide range of autoantibody driven & other inflammation mediated conditions

US Prevalence (2022)



**Key**  
■ Autoantibody driven  
■ Other mechanisms

# Thyroid Eye Disease (TED)



# Each year, TED impacts the lives of ~30k new patients in the US at an average age of ~45<sup>1,2</sup>

## Autoimmune disease associated with proliferation and damaging inflammation of the cell types surrounding the eye

- Disfiguring symptoms that significantly affect QoL: proptosis, double-vision
- Involvement of optic nerve can be sight-threatening, requiring surgery

We estimate up to 2/3 of the 30K new TED patients each year are diagnosed as moderate-to-severe<sup>1,2</sup>

## Pathophysiology driven by autoantibodies that bind to the TSH receptor, which is expressed on cell types surrounding the eye

- Same autoantibody can also cause Graves' hyperthyroidism (GH); up to 95% of TED patients also have GH<sup>3</sup>

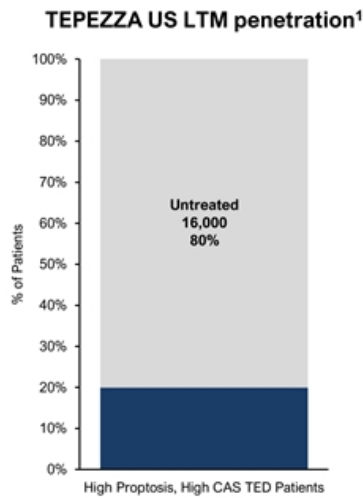


Source: Getty Images

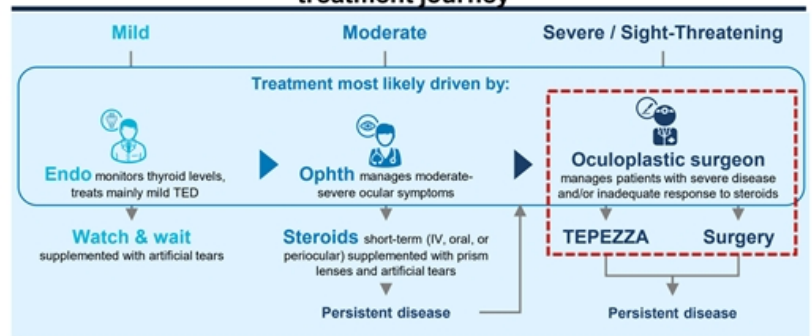
## IL-6 is elevated in patients with TED and a growing body of evidence suggests a role in disease pathogenesis<sup>4</sup>

# Despite an FDA-approved medicine, vast majority of moderate-to-severe TED patients remain untreated

Most TED patients are not receiving TEPEZZA...



...Or only get it relatively late in the treatment journey<sup>2</sup>



## Potential barriers to adoption of TEPEZZA

- Potential safety issues:** risk of potentially permanent hearing loss<sup>3</sup>
- Limited durability:** high relapse rates observed in long-term follow-up<sup>4</sup>
- Inconvenience & complexity:**
  - IV dosing every 3 weeks for a total of 8 infusions<sup>3</sup>
  - Limited access to infusion centers, numerous visits, and time commitment<sup>2,5</sup>
  - Burdensome reimbursement approval process<sup>5</sup>
  - Need for serial audiograms<sup>3</sup>

# FDA label update from July '23 may further impact TEPEZZA's already declining revenues

Despite initial ramp-up, revenues have been declining over last 2 years

Sales by quarter (\$M)<sup>1</sup>

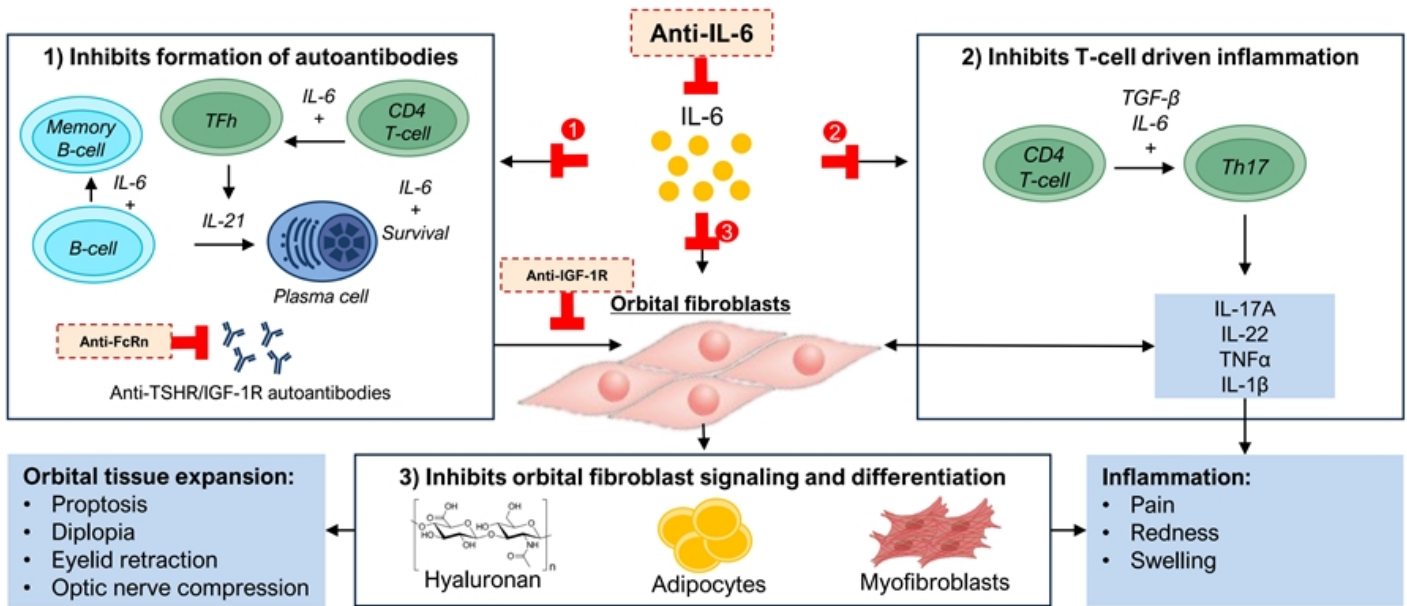


## WARNINGS AND PRECAUTIONS

- **Infusion Reactions:** If an infusion reaction occurs, interrupt or slow the rate of infusion and use appropriate medical management (5.1)
- **Exacerbation of Preexisting Inflammatory Bowel Disease (IBD):** Monitor patients with preexisting IBD for flare of disease; discontinue TEPEZZA if IBD worsens (5.2)
- **Hyperglycemia:** Assess patients for elevated blood glucose and symptoms of hyperglycemia prior to infusion and continue to monitor while on treatment with TEPEZZA. Ensure patients with hyperglycemia or pre-existing diabetes are under appropriate glycemic control before and while receiving TEPEZZA (5.3)
- **Hearing Impairment Including Hearing Loss:** TEPEZZA may cause severe hearing impairment including hearing loss, which in some cases may be permanent. Assess patients' hearing before, during, and after treatment with TEPEZZA and consider the benefit-risk of treatment with patients (5.4)

- 15% of patients reporting hearing impairment across case reports<sup>2</sup>
  - Of these, **45% reported as persistent**
- **427 ear and hearing-related adverse events** captured in the FAERS database, including reports of permanent deafness<sup>3</sup>
- **Ongoing legal actions** filed by patients suffering hearing impairments attributed to teprotumumab across **over 50 lawsuits**<sup>4</sup>
- Hearing impairment likely represents an **on-target consequence of IGF-1 pathway inhibition**<sup>5</sup> (likely just as relevant for SC and oral as IV)

# IL-6 inhibition blocks multiple steps in TED pathogenesis and has potential to play a central and upstream role



# Over 40 publications demonstrate the therapeutic potential of IL-6 pathway inhibition (tocilizumab) in TED

Study Details				Key Endpoints		
First author	Year	Study type	N treated	Proptosis response rate	CAS response rate*	% autoantibody reduction
Pérez-Moreiras	2021	Retro	54	78	89	75
Sánchez-Bilbao	2020	Obs	48	NR	NR	NR
Atienza-Mateo	2018	Retro	29	NR	NR	NR
Pérez-Moreiras	2014	Prosp	18	72	100	76
Pérez-Moreiras	2018	RCT	15	93	60	NS
de la Fuente Bursón	2020	Retro	15	NR	NR	NR
Pereira	2023	Retro	14	NR	NR	NR
Boutzios	2023	Obs	12	NR	NR	84
Pampin-Sánchez	2022	Retro	11	75	73	NR
Moi	2022	Retro	10	CI	80	75
Cortez	2022	Prosp	10	10	100	81
Silkiss	2020	CS	9	CI	56	74
Smith	2021	Retro	9	78	100	54
Bielefeld	2019	Obs	8	NR	NR	NR
Ceballos-Marcias Jose	2020	CS	8	NR	75	41
Benedjal	2020	Retro	7	NR	NR	73
Moás	2022	Obs	7	NR	NR	92
Toro-Tobon	2023	Retro	6	50	NR	NR
de Pablo Gomez	2018	CS	5	NR	60	NR
Ribi	2017	CS	3	33	67	NR
Maldiney	2020	CS	3	67	NR	NR
Stevens	2022	Retro	3	100	67	NR
Russell	2017	CS	2	NR	0	NR

Study Details				Key Endpoints		
First author	Year	Study type	N treated	Proptosis response rate	CAS response rate*	% autoantibody reduction
Sy	2017	CS	2	CI	50	69
Copperman	2019	CS	2	100	0	NR
Coy	2019	CS	2	NR	50	NR
Park	2021	CS	2	100	100	NR
Abeillon-du Payrat	2022	CS	2	100	50	NR
Butnaru	2013	CR	1	NR	100	NR
Gómez Rodríguez	2014	CR	1	NR	100	NR
Bielefeld	2017	CR	1	CI	NR	NR
Canas	2018	CR	1	100	NR	NR
Pascual-Camps	2018	CR	1	NR	NR	NR
Garreta Fontelles	2019	CR	1	NR	NR	93
Mehmet	2020	CR	1	0	NR	NR
Kaplan	2020	CR	1	NR	0	85
Cayon-Blanco	2020	CR	1	NR	100	NR
Tran	2020	CS	1	NR	NR	NR
Ruiz	2021	CR	1	NR	NR	NR
Albrashdi	2022	CR	1	100	NR	NR
Cezara	2022	CR	1	NR	0	NR
Mohamed	2022	CS	1	0	0	NR
Moleiro	2022	CR	1	100	NR	86
Almazrouei	2023	CR	1	NR	NR	NR
Cucuilescu	2023	CR	1	CI	0	NR
Nirmalan	2023	CS	1	NR	NR	NR

	Weighted Mean	72%	78%	74%
Smith 2017 (tepro Phase 2)	71%	69%	N/A	
Douglas 2020 (tepro Phase 3)	83%	59%	N/A	

We believe many of these reports may be underestimating the true efficacy of IL-6 blockade

- 300+ mostly steroid-refractory patients
- Late IL-6 inhibition (>9 months post symptom onset) when disease may have exited active phase
- Exposure to IL-6 inhibition may have been suboptimal (<6 months)

## TOURMALINE

Proptosis response rate is generally defined in the data outlined here as a  $\geq 2$  mm proptosis improvement in the worse eye at baseline without any worsening in the other eye. CAS response rate is generally defined in the data outlined here as a CAS of 0 or 1. Studies referenced in this table represent investigator-led studies and were not designed with the intent of generating evidence for an approval of tocilizumab in TED. The majority of these studies were not designed with power to detect statistical significance. Retro: retrospective, Obs: observational, Prosp: prospective, RCR: randomized controlled trial, CS: case series, CR: case report, NR: not reported, NR: not reported, NS: not significant, CI: clear improvement. Publications available upon request

# TOUR006 has potential to be an optimal first-line TED therapy

## Elements of ideal 1<sup>st</sup> line therapy for TED

**Broad, deep, and durable efficacy**

**Well-tolerated safety profile**

**Physician & patient-friendly experience**

**Resolution of underlying biology**

## Potential improvements to existing therapy

- Meaningful benefits on proptosis, CAS, and diplopia
- Durable response, partially driven by low immunogenicity
- Important improvements on QoL measures

- Well characterized safety profile of IL-6 class across many diseases
- Well tolerated safety profile, manageable with routine monitoring
- Lack of permanent or irreversible side effects

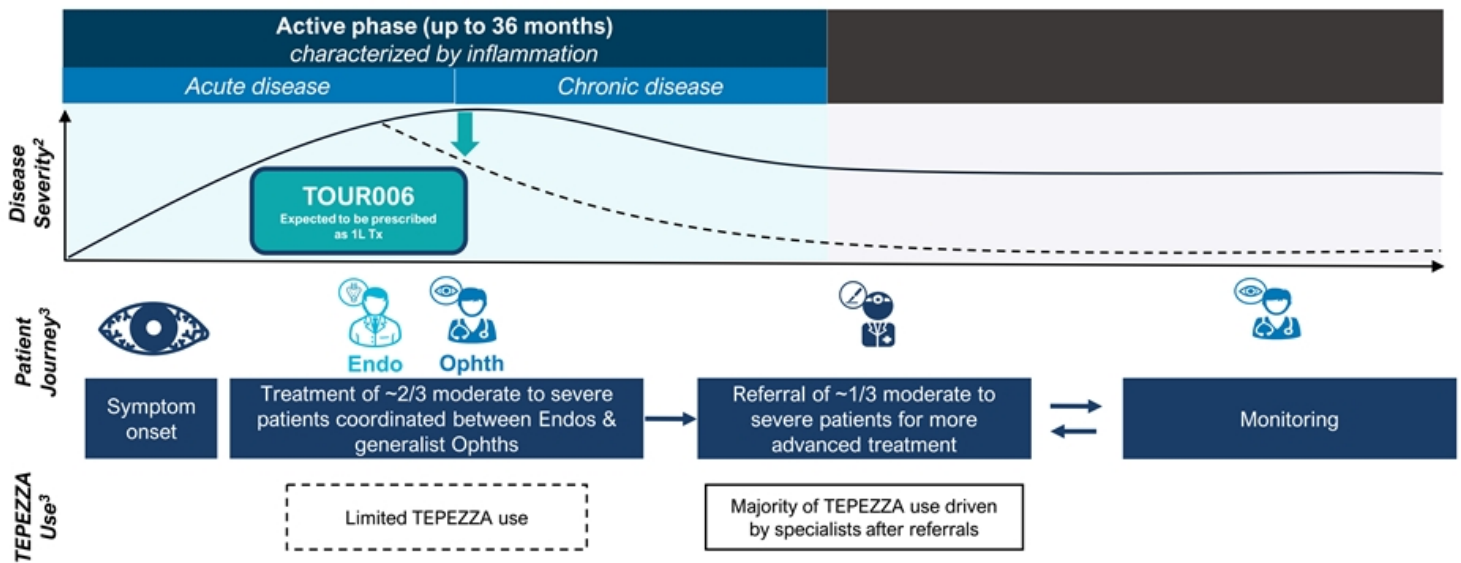
- Subcutaneous, low-volume PFS injections, every 8 weeks
- Finite treatment course for majority of patients (i.e. 6 months)
- Suitable for longer term usage, if indicated

- Mechanistically acts upstream in disease cascade to stop disease
- Clinical data suggests early treatment initiation improves outcomes



# TOUR006 seeks to intervene early, stopping disease progression in active phase that is characterized by inflammation

Moderate-to-severe TED progression & patient journey (up to 20k patients per year in US<sup>1</sup>)

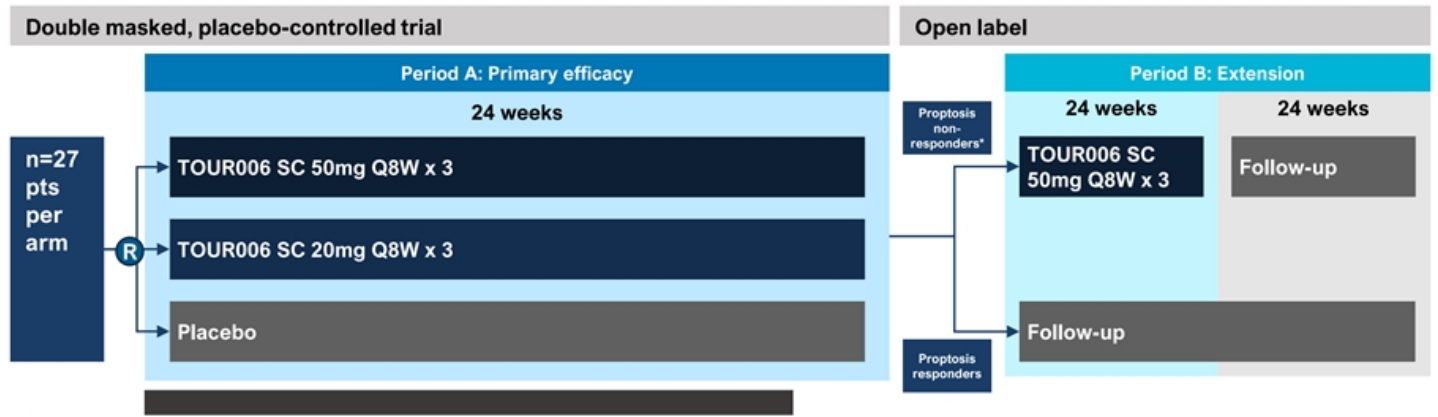


# TOUR006 has potential to offer a well-differentiated profile for first-line TED

MOA	Drug Properties				Dosing interval (days)				
	Drug	Company	Stage	Route of administration	QW	Q2W	Q3W	Q4W	Q8W
IL-6	TOUR006	TOURMALINE	Planned Ph2b start in Q3 '23	SC	56				
	Satralizumab	Roche	Ph3 start expected in 2H '23	SC (with loading dose)	28				
IGF-1R	TEPEZZA (teprotumumab)	HORIZON	Approved	IV	21				
	VRDN-001	VIRIDIAN	Ph3	IV	21				
	Lonigutamab	ACELYRIN	Ph1/2	SC	28				
	Linsitinib	Sling Therapeutics	Ph2/3	Oral	BID				
FcRn	Batoclimab	IMMUNOVANT	Ph3	SC (up to 4mL per administration)	7				
	Efgartigimod	argenx	Ph3 start expected in Q4 '23	IV or SC	7				



# spiriTED dose-ranging Ph2b study in first-line TED



## Study population:

- Moderate-to-severe TED, with proptosis  $\geq 3$ mm above normal (based on race and gender)
- Active phase, with symptom onset  $\leq 12$  months, CAS  $\geq 4$  and positive TSI
- First-line setting, with cap on prior corticosteroid use ( $< 1$ g methylprednisolone or equivalent)

## Primary efficacy endpoint:

- Proptosis response rate at week 20

## Additional endpoints:

- CAS
- Diplopia
- QoL, safety, PK/PD/ADA

# TOUR006: Therapeutic potential to address broad segments of the TED population (and beyond)

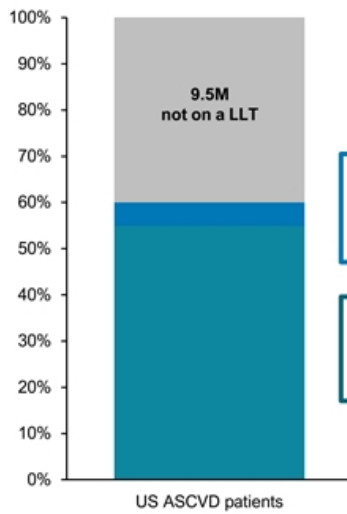
First Line		Post-First Line		Underlying Thyroid Disorder
<b>Active TED (moderate-severe)</b> <ul style="list-style-type: none"> <li>TED treatment-naïve or limited prior treatment (e.g., modest exposure to systemic glucocorticoids)</li> </ul>	<b>High CAS without significant proptosis</b> <ul style="list-style-type: none"> <li>Active inflammation but minimal or no proptosis</li> </ul>	<b>Prior therapy-experienced</b> <ul style="list-style-type: none"> <li>Teprotumumab:                             <ul style="list-style-type: none"> <li>Inadequate response</li> <li>Relapse</li> <li>Unable to tolerate</li> </ul> </li> <li>Other agents (e.g., full glucocorticoid course)</li> </ul>	<b>Extended treatment (beyond initial course)</b> <ul style="list-style-type: none"> <li>Subset of patients that may have inadequate response to TOUR006 after initial 16-week course</li> </ul>	<b>Graves' disease</b> <ul style="list-style-type: none"> <li>Graves without TED: address thyroid disease</li> <li>Pre-TED: prevent TED</li> <li>Early/mild stages of TED: stabilize or reverse</li> </ul>
<i>Evaluate through Phase 2b trial</i>	<i>Evaluate through TED basket trial</i>		<i>Evaluate through Phase 2b trial</i>	<i>Evaluate through other trial(s)</i>

## Key TED milestones

Indication	Milestone	Expected timing	Status
TED	Gain FDA alignment on proposed TED program	Q2 2023	<input checked="" type="checkbox"/>
	File TED IND	Mid-2023	<input checked="" type="checkbox"/>
	Receive TED IND FDA clearance	August 2023	<input checked="" type="checkbox"/>
	Initiate Phase 2b TED trial	Q3 2023	<input type="checkbox"/>
	Initiate TED basket trial	Early 2024	<input type="checkbox"/>
	Report Phase 2b TED trial topline results	H1 2025	<input type="checkbox"/>

# Atherosclerotic Cardiovascular Disease (ASCVD)

# ASCVD continues to be underserved despite the wide availability of lipid lowering therapies



Other lipid-lowering therapies (LLTs) (e.g. PCSK9 class) with **<5% usage** in US ASCVD patients<sup>3</sup>

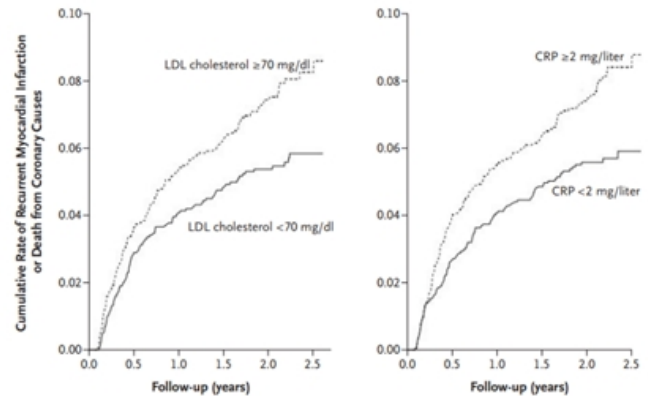
~55% of US patients get put on statins, of which **only 50% have an adequate response**<sup>3,4</sup>

- Dyslipidemia
- Diabetes
- Obesity
- Hypertension
- Chronic kidney disease

• **Persistent inflammation**

# Decades of research have indicated elevated IL-6 driven inflammation as a predictor of major CV events

- High levels of CRP are a known risk factor for ASCVD, nearly tripling risk of occurrence of MACE in one study<sup>1</sup>
- CRP levels  $\geq 2.0$  mg/L is listed as a risk-enhancing factor alongside elevated LDL-C and other well-known risk factors by the ACC & AHA<sup>2</sup>
- Chronic inflammatory conditions associated with elevated CRP such as RA are also included along with primary hypercholesterolemia and metabolic disorders as potential risk factors<sup>2</sup>
- Multiple large cardiovascular outcome studies have demonstrated reductions in CRP were associated with improved outcomes and has been a powerful predictor for therapeutic benefit<sup>3</sup>



# Analysis of CANTOS data implicates IL-6 as a key risk factor for ASCVD

Canakinumab, an anti-IL-1 $\beta$  antibody that partially decreases IL-6 levels, demonstrated in a Phase 3 CV outcomes trial (CANTOS) greater IL-6 and hsCRP reductions are associated with greater CV benefit

**CANTOS primary endpoint<sup>1</sup>**

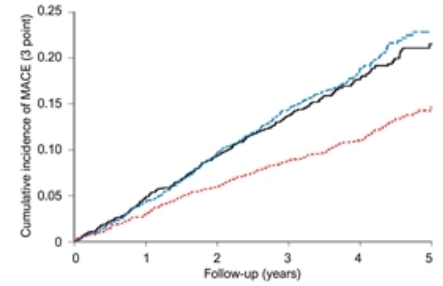
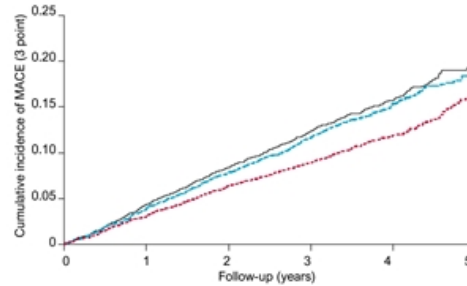
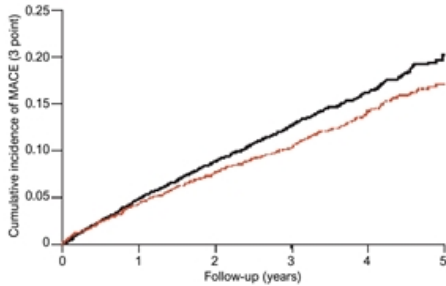
	HR	(95% CI)	p
Placebo	1	(ref)	(ref)
Canakinumab, 150 mg	<b>0.85</b>	(0.74-0.98)	P=0.021

**CANTOS stratified by hsCRP reductions<sup>2</sup>**

	HR	(95% CI)	p
Placebo	1	(ref)	(ref)
On-treatment hsCRP > 2.0 mg/L	0.95	(0.84-1.09)	0.48
On-treatment hsCRP < 2.0 mg/L	<b>0.75</b>	(0.66-0.85)	<0.0001

**CANTOS stratified by IL-6 reductions<sup>3</sup>**

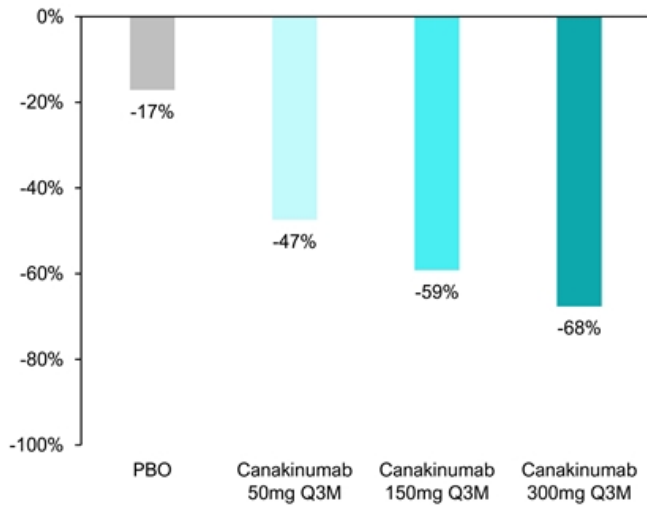
	HR	(95% CI)	p
Placebo	1	(ref)	(ref)
On-treatment IL-6 > 1.65 ng/L	1.06	(0.90-1.25)	0.49
On-treatment IL-6 < 1.65 ng/L	<b>0.64</b>	(0.54-0.77)	<0.0001



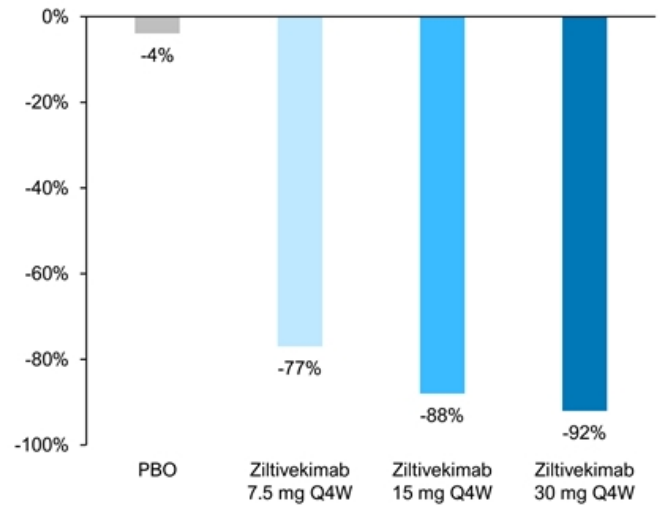
Hazard ratio improvement

# Ziltivekimab, an anti-IL-6 antibody developed by Corvidia, produced deeper CRP reductions than canakinumab

Canakinumab only achieved 59-68% median CRP reduction at higher doses at week 12<sup>1</sup>



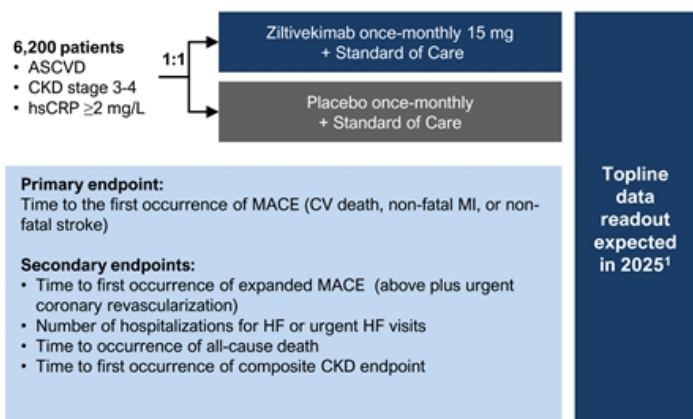
Primary endpoint of Phase 2b RESCUE study: 92% reduction in median CRP at week 12<sup>2</sup>



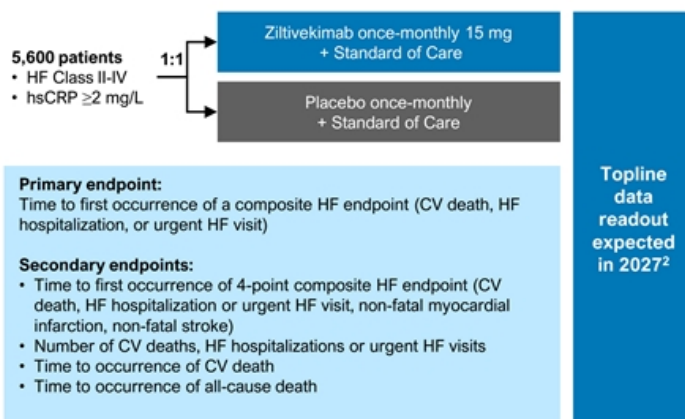


# Novo studying ziltivekimab in two Phase 3 CV outcomes trials in ASCVD patients with kidney disease and heart failure

## ZEUS trial design in ASCVD with CKD<sup>1</sup>



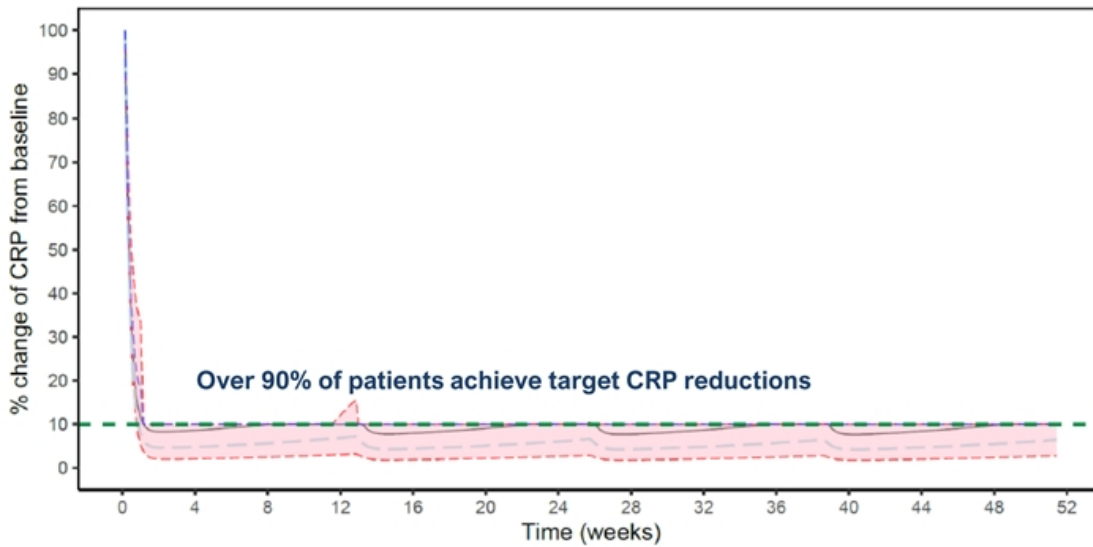
## HERMES trial design in heart failure (HF)<sup>2</sup>



**Potential opportunity for TOUR006 to enter market with a less frequently administered product as a fast follower**

# PK/PD modeling for TOUR006 supports potential for quarterly administration

Simulations with 50mg every 90 days with CRP >2mg/L & <10mg/L



Black straight line is the median

Red dotted lines are the 5<sup>th</sup> and the 95<sup>th</sup> percentiles

Blue dotted lines are the 25<sup>th</sup> and 75<sup>th</sup> percentiles

Green dashed line 90% decline of the CRP from baseline

*Note: To mitigate against ceiling effects from CRP levels entering into the normal range, any simulated patient with CRP attaining a value < 2 mg/L was considered to have achieved a 90% decrease from baseline CRP*

# Planned Phase 2 CV trial\*



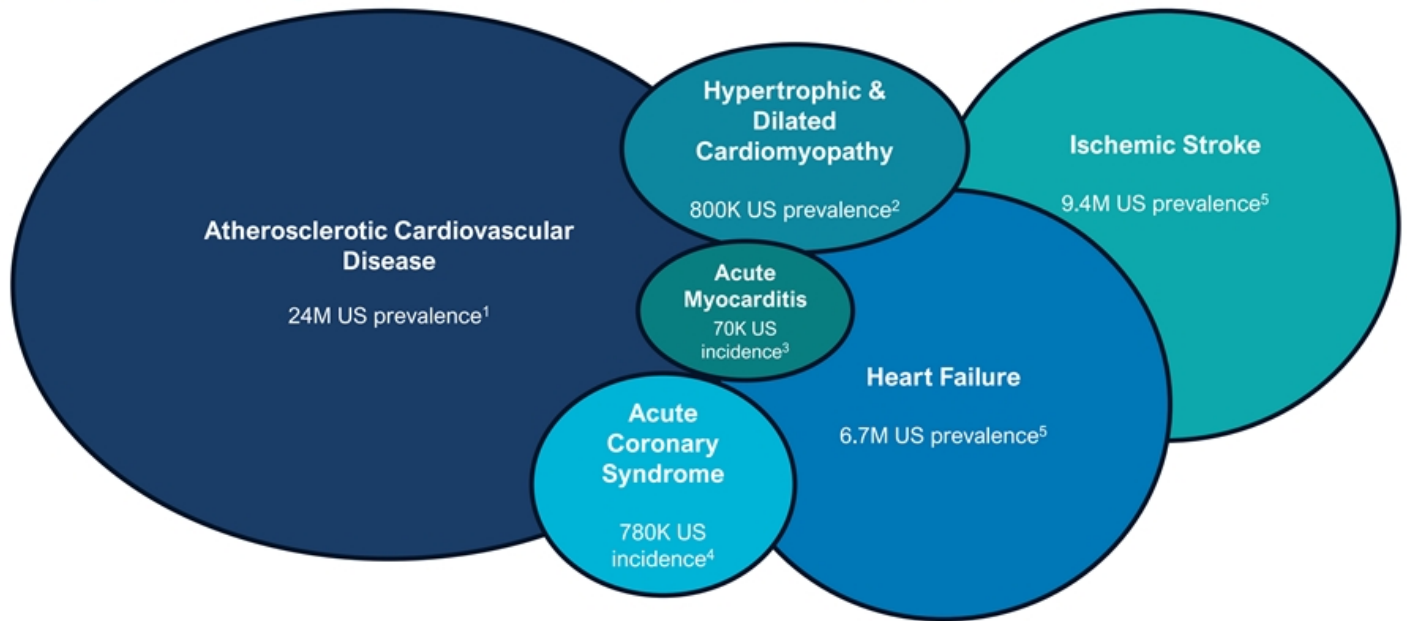
**Study population:**

- Study population similar to RESCUE trial
- Exclude patients at higher risk for safety complications (e.g., immunocompromised patients)

**Key endpoints:**

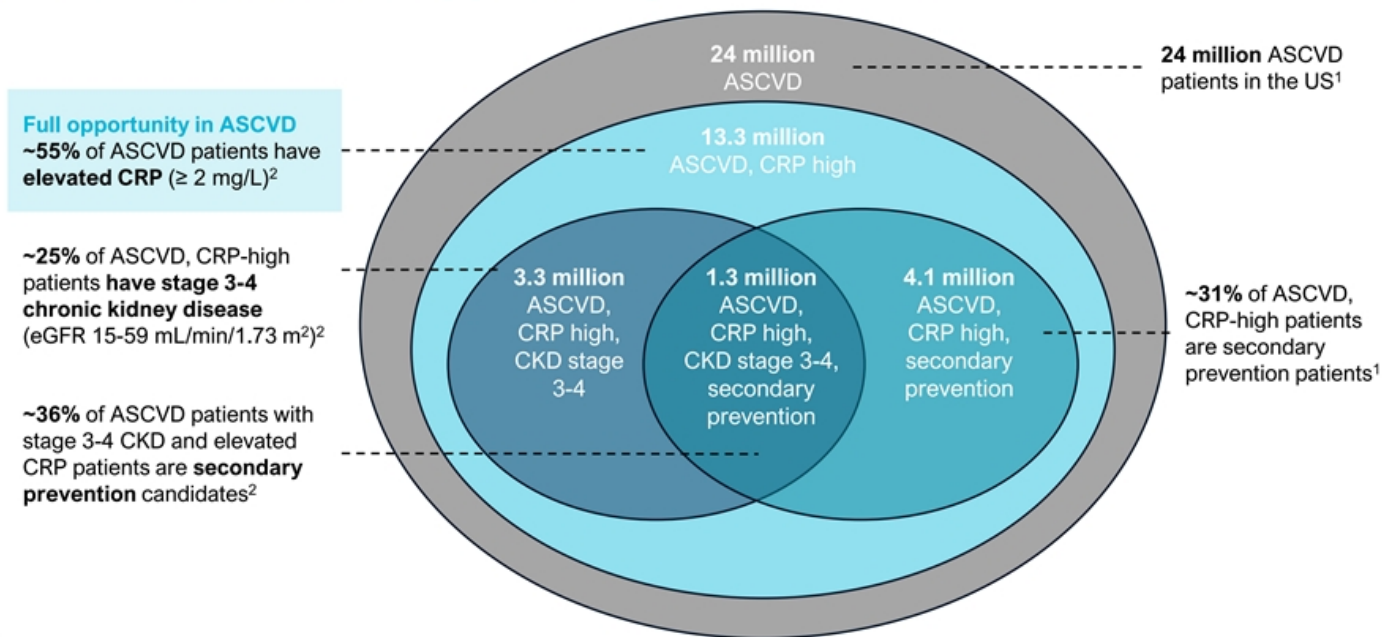
- PD: hsCRP and other biomarkers
- PK, ADA
- Safety

# IL-6 potentially implicated across cardiovascular disorders



*Illustrative, non-exhaustive and non-mutually exclusive; bubbles sizes not to scale*

# Illustrative ASCVD opportunity in US alone



## Key ASCVD milestones

Indication	Milestone	Expected timing	Status
ASCVD	Gain FDA alignment on proposed CV program	Q4 2023	<input type="checkbox"/>
	File ASCVD IND	H1 2024	<input type="checkbox"/>
	Receive ASCVD IND clearance	2024	<input type="checkbox"/>
	Initiate Phase 2 ASCVD trial	2024	<input type="checkbox"/>
	Report topline Phase 2 ASCVD trial results	2025	<input type="checkbox"/>

# Key Business Items

# TOURMALINE

**Mission Statement: Developing transformative medicines that dramatically improve the lives of patients with life-altering immune diseases**

## Our Core Values and Corporate Behaviors

### We have passion for our mission

- Understand and strive to address patient needs
- "All hands on deck" mindset
- Follow through to the last mile and beyond

### We believe respect and inclusion are core to the success of our team

- Operate in a respectful, transparent, and honest way
- Demonstrate that the diverse experience and perspective of team members is valued
- Collaborate with kindness

### We overcome obstacles to deliver results for patients

- Focus on the problem, not on individuals or groups
- Think creatively and act quickly to apply solutions
- Draw on strengths of the whole team

### We push the envelope

- Challenge the status quo
- Take well-informed risks
- Find inspiration and opportunity within and beyond Tourmaline



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## Terms of TOUR006 license with Pfizer

In May 2022, Tourmaline obtained an exclusive global license to TOUR006 (formerly PF-04236921) from Pfizer in exchange for:

- \$5M upfront payment and 15% equity of Tourmaline
- Milestones:
  - Up to \$128M in development and regulatory milestones
  - Up to \$525M in sales-based milestones
- Low double-digit (less than 15%) royalties on net sales, subject to specified royalty reductions
- Additional payments due upon out-license/divestiture of TOUR006 in major market
- Right of first negotiation for Pfizer should Tourmaline seek to out-license or partner in the US

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# TOUR006 regulatory exclusivity and intellectual property

## Regulatory Exclusivity

- In the US, we expect to rely on 12 years' data exclusivity for biologics
  - Regulatory counsel has confirmed this is a reasonable expectation

## Patent Protection

- We have filed 5 new patent applications on TOUR006
  - Incorporating claims on:
    - Indication-specific methods of use
    - Dosing regimens
  - If issued, will expire in 2043 (or later)
- Additional patent applications in process
- Pfizer has abandoned all previous Pfizer patents/applications relating to TOUR006
- No freedom to operate issues identified

## Reverse merger transaction summary

<b>Overview</b>	<ul style="list-style-type: none"> <li>• Talaris Therapeutics, Inc. (NASDAQ: TALS) to acquire 100% of outstanding equity interests of Tourmaline Bio, Inc. structured as a traditional reverse merger</li> <li>• Surviving entity name / proposed ticker: Tourmaline Bio (TRML)</li> <li>• Issuer of shares in private placement: Tourmaline</li> </ul>
<b>Transaction Summary</b>	<ul style="list-style-type: none"> <li>• \$389.7M pro forma value of combined company               <ul style="list-style-type: none"> <li>– \$230M value of Tourmaline</li> <li>– \$84.7M value of Talaris including net of up to \$64.6M dividend/equity award cash payout<sup>1</sup></li> <li>– \$75M private placement</li> </ul> </li> <li>• ~\$210M pro forma cash balance for combined company estimated at close excluding dividend</li> <li>• Pro forma ownership split: ~59.0% Tourmaline, ~21.7% Talaris, ~19.3% private placement</li> <li>• Private placement syndicate includes Acuta Capital Partners, Affinity Asset Advisors, Braidwell LP, Cowen Healthcare Investments, Deep Track Capital, Great Point Partners, LLC, KVP Capital, Logos Capital, Paradigm BioCapital, Qiming Venture Partners USA, RA Capital Management, LP, StemPoint Capital LP, TCGX, Vivo Capital, and other undisclosed investors</li> </ul>
<b>Use of Proceeds</b>	<ul style="list-style-type: none"> <li>• Expected to fund Tourmaline through 2026 and provide sufficient capital for key clinical programs including Phase 2b TED study, Phase 2 TED basket study, and Phase 2 CV study</li> </ul>
<b>Projected Timing</b>	<ul style="list-style-type: none"> <li>• Closing expected Q4 2023 – shareholder vote scheduled October 17, 2023</li> </ul>

## Key highlights



An IL-6 renaissance is underway: new insights emerging about a broad range of indications where IL-6 may be clinically validated

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TOUR006 offers potential for low volume, infrequent subcutaneous administration

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We are rapidly advancing TOUR006 into mid/late-stage development

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Our team has extensive experience developing and commercializing antibodies for orphan and autoimmune diseases

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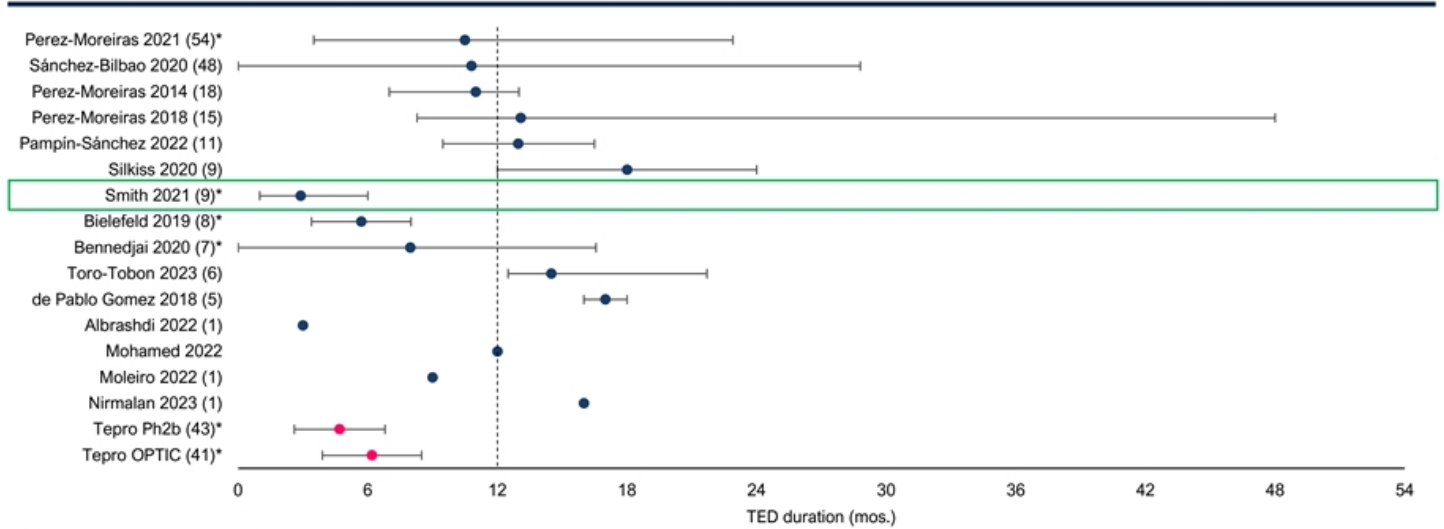


Cash runway expected to fund development through 2026\*

# Appendix

# TED patients had mostly long disease durations before starting tocilizumab, likely impacting efficacy of IL-6 blockade

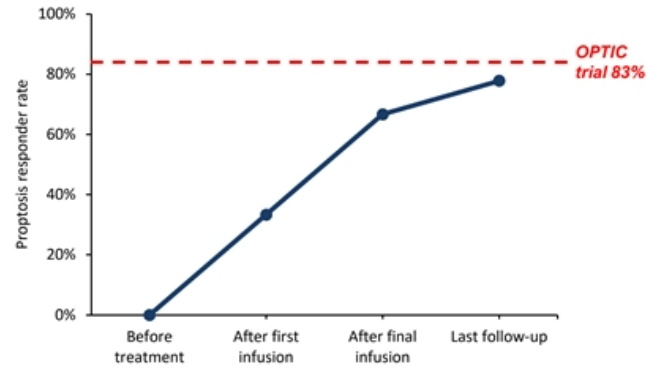
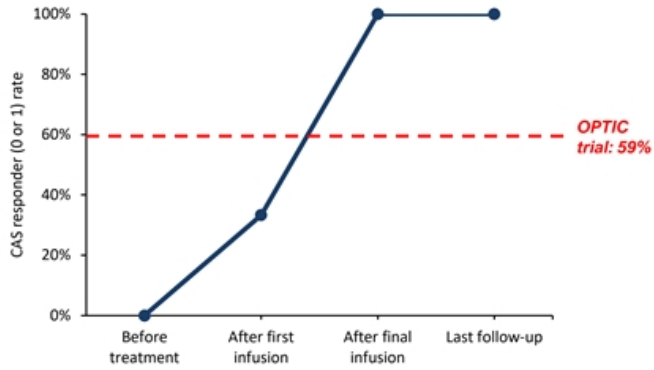
## Median (IQR) duration of TED symptoms at time of enrollment (n)








# Recent case series demonstrate IL-6 inhibition's potential in first-line TED patients

## Investigator-led retrospective analysis using tocilizumab

- 9 subjects included in analysis
- Average CAS: 6 (out of 7)
- Treatment with tocilizumab 8mg/kg monthly
- Mean time from symptom onset to first treatment: 2.89 months
- Mean number of infusions: 4.2
- Median change in autoantibody levels from baseline: 61% decline



# Treatment of ASCVD is focused on preventing major events by reducing risk factors, yet inflammation remains unaddressed

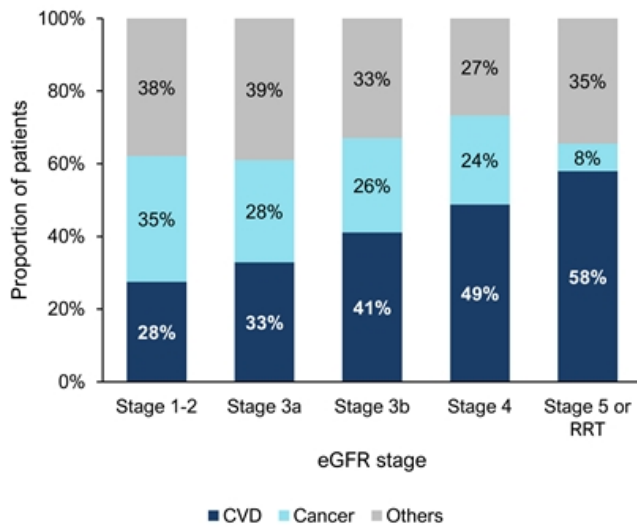
ACC/AHA risk factor	Dyslipidemia (high LDL-C)	Metabolic syndrome (diabetes & obesity)	Hypertension (high blood pressure)	Inflammation (high CRP)
<b>Legacy treatments</b>				—
<b>New therapies</b>			—	<b>LODOCO</b> (coichicine) tablets
<b>New mechanisms in development</b>	Lp(a) inhibitors/ASOs ANGPTL3 ASOs Oral PCSK9 inhibitors CETP inhibitors	Oral, non peptide GLP-1 agonists GGG triple agonist amylin agonists	Angiotensinogen ASOs Aldosterone synthase inhibitor Endothelin receptor antagonist NPR1 agonist	<b>IL-6 mAbs</b>

**Inflammation as a risk factor remains underexplored**

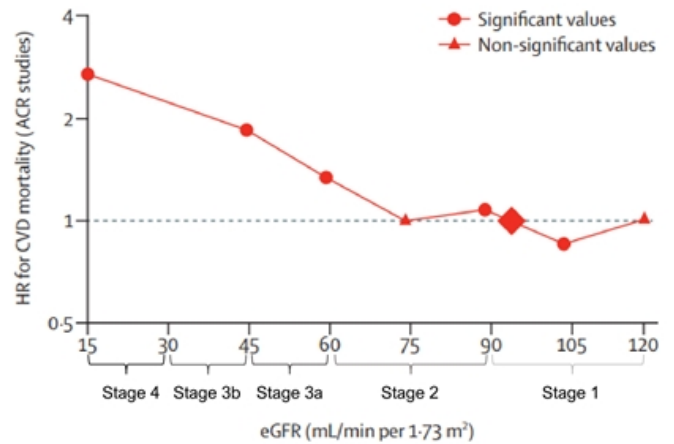


# ASCVD patients with chronic kidney disease (CKD), an inflammatory condition, have elevated cardiovascular risk

**Risk of CV deaths increases as CKD worsens<sup>1</sup>**

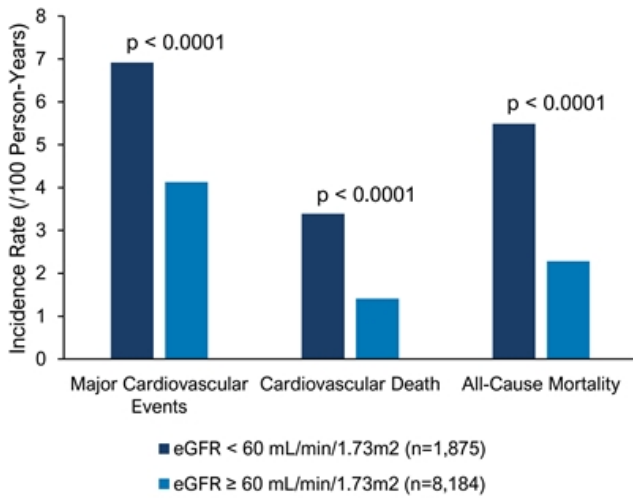


**Double and triple risk of CV mortality in Stage 3 and 4 CKD patients<sup>1</sup>**

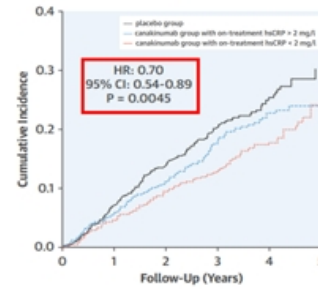


# Patients in CANTOS with CKD had higher MACE risk and demonstrated amplified benefit from inflammation reduction

## Higher MACE rates observed in CKD patients<sup>1</sup>



## MACE benefit from CRP reductions in CANTOS CKD patients<sup>1</sup>



### Relative risk reduction in patients who achieved hsCRP < 2 mg/L at 3 months vs PBO

	CANTOS, all patients <sup>2</sup>	CANTOS, CKD patients <sup>1</sup>
MACE	25%	30%
MACE+	26%	32%
CV Mortality	31%	39%

**TOURMALINE**