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# **COSTS AND HEALTHCARE RESOURCE UTILIZATION IN TRANSTHYRETIN AMYLOID CARDIOMYOPATHY EXCEEDS THAT OF NON-AMYLOID HEART FAILURE**

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# DISCLOSURE OF RELEVANT FINANCIAL RELATIONSHIPS WITH INDUSTRY AND ACKNOWLEDGMENTS

**Sandesh Dev** has participated in advisory boards (uncompensated) for Pfizer and BridgeBio Pharma, Inc., has received grants from Pfizer, and has no consultancy fees to disclose.

**Justin Grodin** is a researcher for Texas Health Resources Clinical Scholarship, BridgeBio Pharma, Inc., Pfizer, and NHLBI R01HL160892, and a consultant, advisor, and speaker for Pfizer, BridgeBio Pharma, Inc., AstraZeneca, Intellia, Tenax Therapeutics, and Alexion.

**Ahmad Masri** is a researcher for Pfizer, Ionis, Attralus, and Cytokinetics, and a consultant, advisor, and speaker for Cytokinetics, BMS, BridgeBio Pharma, Inc., Pfizer, Ionis, Lexicon, Attralus, Alnylam, Haya, Alexion, Akros, Prothena, BioMarin, AstraZeneca, and Tenaya.

**Richard Wright** is a consultant, advisor, and speaker for Alnylam, Amgen, AstraZeneca, BMS, Boehringer Ingelheim, BridgeBio Pharma, Inc., Cytokinetics, Lexicon, Lilly, Myocardia, and Novartis.

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# INTRODUCTION

- ATTR-CM is a progressive, fatal condition characterized by worsening HF, exercise intolerance, cardiac arrhythmias, aortic stenosis, and extracardiac manifestations<sup>1-5</sup>
- Although the prevalence of ATTR-CM is not well characterized due to high rates of misdiagnosis and delayed diagnosis,<sup>1,4,6</sup> it has been estimated that up to 150,000 people in the US have HF caused by ATTR-CM<sup>3</sup>
- The prognosis for patients with ATTR-CM is poor, with many studies reporting a median survival time of approximately 3-5 years, depending on the disease type and stage at the time of diagnosis<sup>3-7</sup>
- Previous analyses of patients in Europe and Asia have shown that ATTR-CM is associated with a substantial burden to healthcare systems<sup>6-8</sup>; however, there are limited data available to understand HCRU for patients with ATTR-CM in the US
- Additionally, it is important to understand how the burden of disease among patients with ATTR-CM compares to those with non-amyloid HF



## OBJECTIVE:

To compare all-cause and CV-related HCRU and costs between patients with ATTR-CM and patients with non-amyloid HF in the US using real-world medical and pharmacy claims data

ATTR-CM, transthyretin amyloid cardiomyopathy; CV, cardiovascular; HCRU, healthcare resource utilization; HF, heart failure; US, United States.

1. Rozenbaum MH, et al. *Cardiol Ther*. 2021;10(1):141-159. 2. Maurer MS, et al. *Circ Heart Fail*. 2019;12(9):e0067075. 3. Ruberg FL, et al. *JAMA*. 2024;331(9):778-791. 4. Jain A, Zahra F. StatPearls [Internet]. 2024. 5. Gillmore JD, et al. *Circulation*. 2016;133(24):2404-2412. 6. Lane T, et al. *Circulation*. 2019;140(1):16-26. 7. Jang SC, et al. *Orphanet J Rare Dis*. 2022;17(1):262. 8. Lauppe R, et al. *ESC Heart Fail*. 2022;9(3):1636-1642.

# METHODS

**Data source:** Optum® Clinformatics® Data Mart database (Jan 2016–Sept 2023)

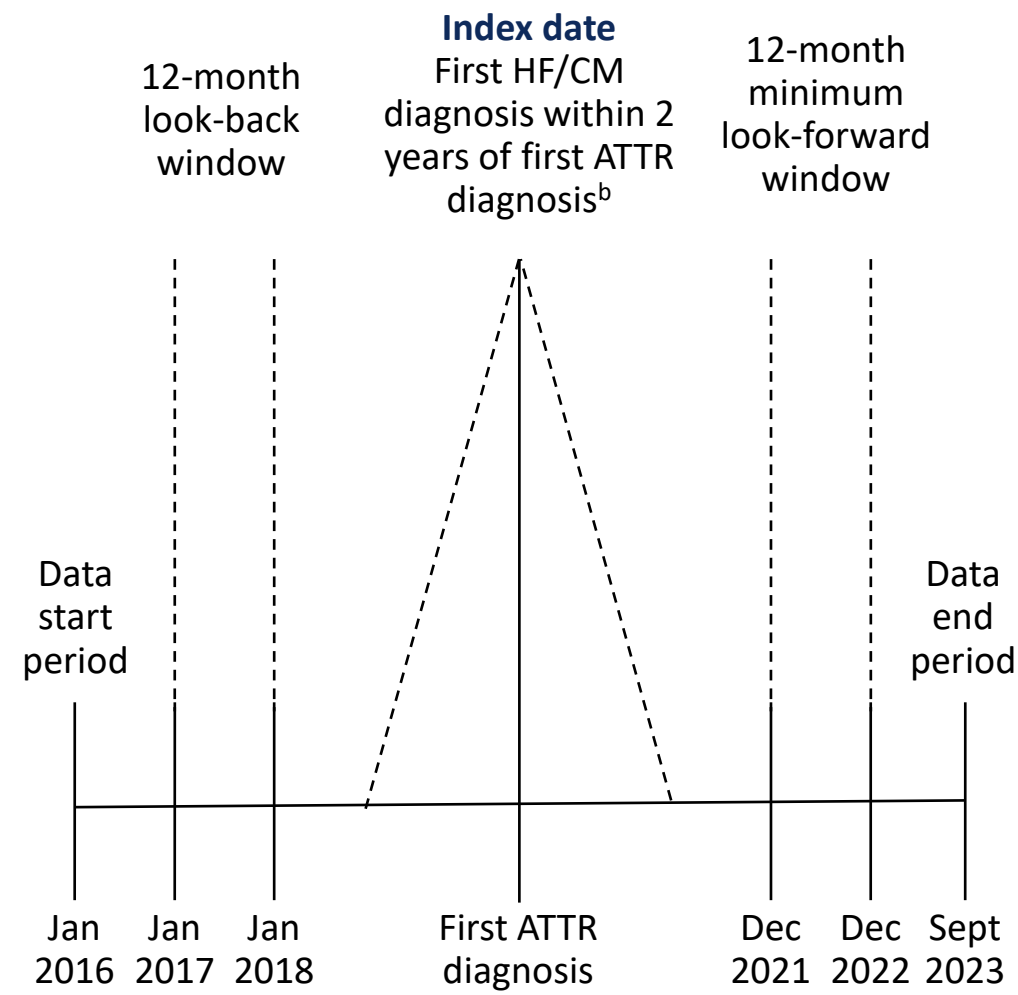
**Inclusion criteria:**

- Diagnosis of ATTR<sup>a</sup> on any claim (Jan 2017–Dec 2022)
- Diagnosis of HF and/or CM<sup>a</sup> within 2 years of first ATTR diagnosis
- Minimum 2 years of continuous enrollment (with minimum 3-month look-back and 12-month look-forward from index<sup>b</sup>)
- Excluded if diagnosed with AL amyloidosis or MM, or received chemotherapy or heart, liver, or kidney transplant (2016–2022)

**Comparison group:** Non-amyloid HF cohort (diagnosis of HF<sup>a</sup> with no evidence of ATTR)

**Outcomes:** Baseline demographics, procedures of interest, all-cause and CV-related hospitalizations<sup>c</sup> and LOS, and hospitalization-related inpatient costs (2024 USD)

**Statistical analysis:** Patients with ATTR-CM were matched using 1:1 PSM<sup>d</sup> to patients with non-amyloid HF. Hospitalizations/costs were compared using 2-tailed t tests and Mann-Whitney U tests



<sup>a</sup>Based on ICD-10-CM codes. <sup>b</sup>First HF/CM diagnosis after first ATTR diagnosis or last HF/CM diagnosis before first ATTR diagnosis if no post-ATTR HF/CM diagnosis. <sup>c</sup>CV-related hospitalizations were defined as inpatient admissions during which the patient received a CV diagnosis. <sup>d</sup>PSM methods included 1:1 nearest neighbor matching with a caliper width of 0.2 SD of the logit scores, and was based on the following covariates: age, race, US state, sex, and index diagnosis year.  
AL, light chain; ATTR-CM, transthyretin amyloid cardiomyopathy; CM, cardiomyopathy; CV, cardiovascular; HF, heart failure; ICD-10-CM, International Classification of Diseases, Tenth Revision, Clinical Modification; LOS, length of stay; MM, multiple myeloma; PSM, propensity score matching; SD, standard deviation; USD, United States dollars.

# BASELINE DEMOGRAPHICS AND PROCEDURES OF INTEREST

## Baseline demographics

- 4,966 patients with ATTR-CM and 861,507 patients with non-amyloid HF were identified
- Each matched cohort included 4,571 patients with well-balanced baseline demographics

Characteristic	ATTR-CM (N = 4,571)	Non-amyloid HF (N = 4,571)
Age, years, mean (SD)	75.3 (9.1)	75.5 (8.8)
Sex, male, n (%)	2,570 (56.2)	2,562 (56.0)
Race, n (%)		
Asian	88 (1.9)	93 (2.0)
Black	1,023 (22.4)	729 (15.9)
White	2,949 (64.5)	3,306 (72.3)
Unknown/None <sup>a</sup>	511 (11.2)	443 (9.7)
Ethnicity, n (%)		
Hispanic	365 (8.0)	408 (8.9)
Non-Hispanic	2,243 (49.1)	2,162 (47.3)
Unknown/None <sup>a</sup>	1,963 (42.9)	2,001 (43.8)
Follow-up time, years, mean (SD)	2.9 (1.1)	3.0 (1.2)

<sup>a</sup>Unknown and none are separate categories but have been combined for presentation.  
ATTR-CM, transthyretin amyloid cardiomyopathy; CV, cardiovascular; EGD, esophagogastroduodenoscopy; HF, heart failure; ICD, implantable cardioverter-defibrillator; IV, intravenous; SD, standard deviation.

## Procedures of interest (1-year look-forward from index)

- The most common procedures in both cohorts included cardiac catheterization and coronary angiography, IV diuretics, endoscopy, and colonoscopy

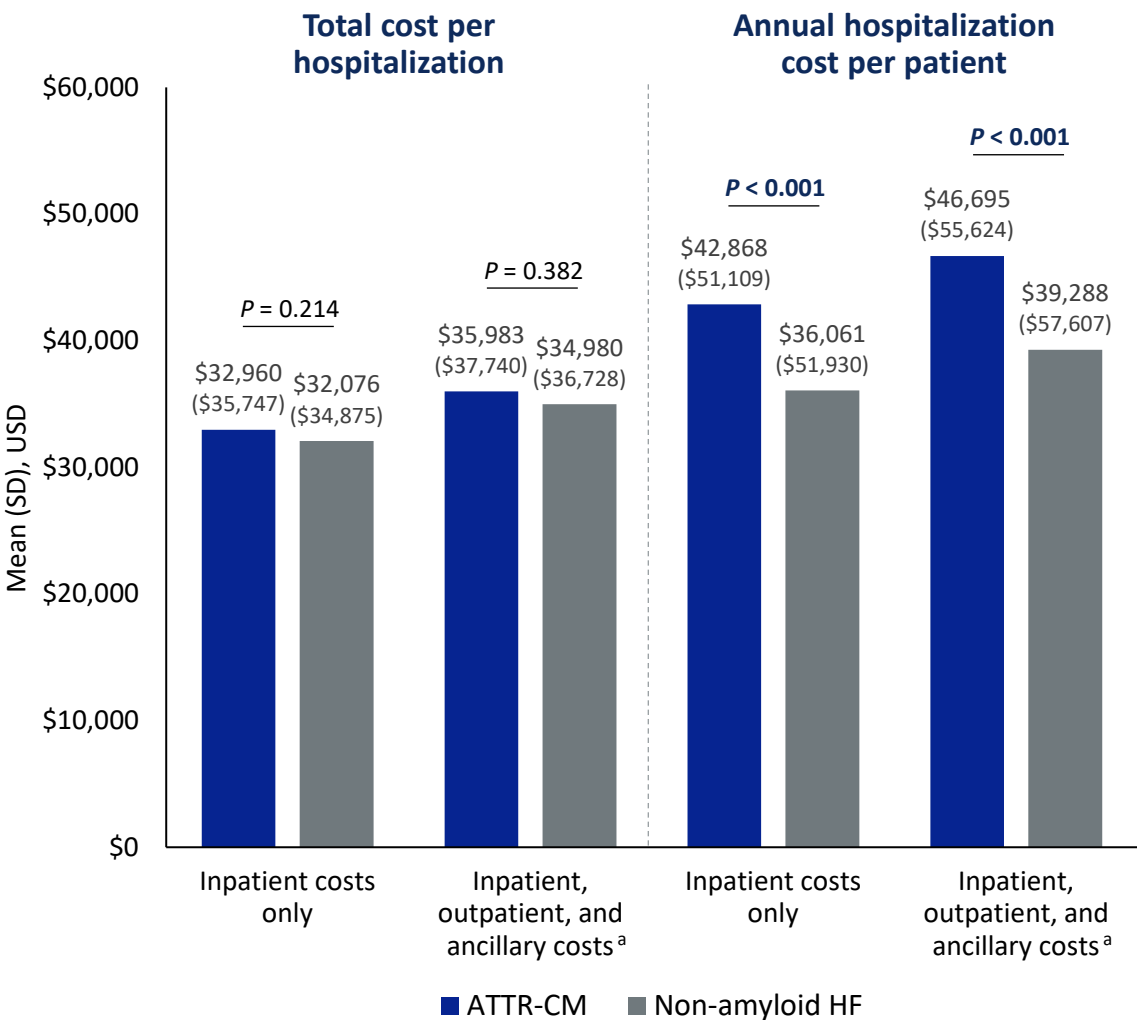
Body system	Procedures, n (% of total cohort)	ATTR-CM (N = 4,571)	Non-amyloid HF (N = 4,571)
N/A	Any procedure of interest	2,412 (52.8)	2,077 (45.4)
CV	Cardioversion	287 (6.3)	166 (3.6)
	Pacemaker + ICD implantation	274 (6.0)	216 (4.7)
	Cardiac catheterization + coronary angiography	1,092 (23.9)	819 (17.9)
	Cardiac catheter ablation	131 (2.9)	78 (1.7)
	IV diuretics	493 (10.8)	372 (8.1)
	Cardiac biopsy	92 (2.0)	0 (0.0)
Digestive	Colonoscopy	392 (8.6)	414 (9.1)
	Upper gastrointestinal endoscopy – EGD	482 (10.5)	473 (10.3)
	Biopsy of liver	16 (0.4)	9 (0.2)
	Abdominal paracentesis	33 (0.7)	24 (0.5)
Eye	Cataract surgery	196 (4.3)	191 (4.2)
Hemic and lymphatic	Bone marrow biopsy	18 (0.4)	11 (0.2)
Musculo-skeletal	Carpal tunnel release	60 (1.3)	22 (0.5)
	Trigger finger/biceps tendons	188 (4.1)	128 (2.8)
	Achilles tendon	2 (0.0)	1 (0.0)
	Laminectomy	35 (0.8)	28 (0.6)
Nervous	Nerve conduction studies + electromyography	240 (5.3)	159 (3.5)

# ALL-CAUSE HOSPITALIZATIONS AND ASSOCIATED COSTS

Compared with the non-amyloid HF cohort, the ATTR-CM cohort had:

- more hospitalizations
- longer LOS per hospitalization
- more days hospitalized annually per patient
- higher annual hospitalization costs per patient

	ATTR-CM (N = 4,571)	Non-amyloid HF (N = 4,571)	P-value
Patients with a hospitalization, n (%)	3,451 (75.5)	3,001 (65.7)	0.433
Total hospitalizations, n	11,255	8,150	< 0.001
Total hospitalizations per patient (95% CI)	3.3 (3.1, 3.4)	2.7 (2.6, 2.9)	< 0.001
LOS per hospitalization, days Mean (SD) Median (95% CI)	8.0 (11.7) 6.0 (5.7, 6.3)	7.5 (7.2) 5.0 (4.8, 5.3)	< 0.001
Days hospitalized annually per patient Mean (SD) Median (95% CI)	10.6 (15.8) 5.9 (5.6, 6.2)	8.4 (12.3) 4.5 (4.3, 4.7)	< 0.001



<sup>a</sup>Ancillary costs include those for durable medical equipment, drugs administered, home health/hospice visits, services/supplies, and transportation incurred during hospitalizations. ATTR-CM, transthyretin amyloid cardiomyopathy; CI, confidence interval; HF, heart failure; LOS, length of stay; SD, standard deviation; USD, United States dollars.

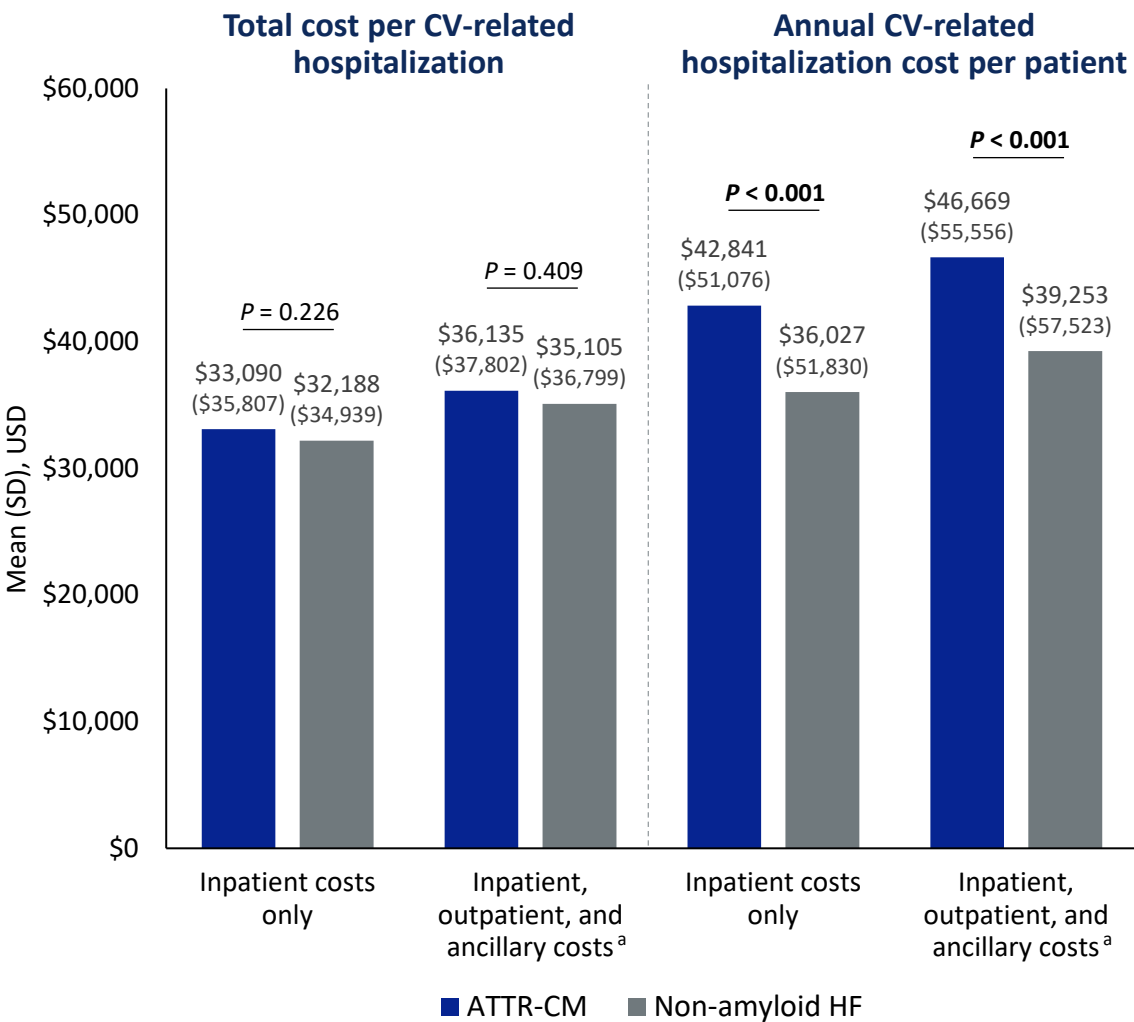
# CV-RELATED HOSPITALIZATIONS AND ASSOCIATED COSTS

Compared with the non-amyloid HF cohort, the ATTR-CM cohort had:

- more CV-related hospitalizations
- longer LOS per CV-related hospitalization
- more CV-related days hospitalized annually per patient
- higher annual CV-related hospitalization costs per patient

Commercial patients had higher mean total costs per CV-related hospitalization than Medicare patients in both cohorts (ATTR-CM cohort: \$47,645 vs \$35,975,  $P < 0.001$ )

	ATTR-CM (N = 4,571)	Non-amyloid HF (N = 4,571)	P-value
Patients with a hospitalization, n (%)	3,440 (75.3)	2,991 (65.4)	0.347
Total hospitalizations, n	11,170	8,085	< 0.001
Total hospitalizations per patient (95% CI)	3.3 (3.1, 3.4)	2.7 (2.6, 2.8)	< 0.001
LOS per hospitalization, days			
Mean (SD)	8.0 (11.7)	7.5 (7.2)	<0.001
Median (95% CI)	6.0 (5.7, 6.3)	6.0 (5.7, 6.3)	
Days hospitalized annually per patient			
Mean (SD)	10.5 (15.6)	8.3 (12.1)	< 0.001
Median (95% CI)	5.9 (5.6, 6.2)	4.5 (4.3, 4.8)	



<sup>a</sup>Ancillary costs include those for durable medical equipment, drugs administered, home health/hospice visits, services/supplies, and transportation incurred during hospitalizations. ATTR-CM, transthyretin amyloid cardiomyopathy; CI, confidence interval; CV, cardiovascular; HF, heart failure; LOS, length of stay; SD, standard deviation; USD, United States dollars.

# CONCLUSIONS



The results of this study suggest that ATTR-CM is associated with an equal or greater per-patient burden on the US healthcare system than non-amyloid HF in terms of hospitalizations and associated costs, with the vast majority (>99%) of hospitalizations being CV related in both groups



Although the management of ATTR-CM has improved with the introduction of the first disease-modifying therapy in 2019,<sup>1,2</sup> the present findings indicate that patients with ATTR-CM have a higher burden of disease and resource utilization than patients with non-amyloid HF that may be addressed by implementing novel disease-specific management strategies



Further research is also warranted to determine whether timely diagnosis and early treatment with current and emerging therapies reduces disease burden and resource utilization due to ATTR-CM at the health system level