Survey Results from Italy, the US, UK, and France: Anxiety in Patients Using Injectable On-Demand **Treatments for Hereditary Angioedema Attacks**

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Background

Studies report a high prevalence of anxiety among patients with hereditary angioedema (HAE). Currently, all approved on-demand treatments for managing these attacks require parenteral administration, which can be painful and challenging to administer and may contribute to treatment-related anxiety¹⁻³

Objective

This study aimed to quantify levels of anxiety associated with the use of injectable on-demand therapies

Methods

- Patients with Type 1 or Type 2 HAE from Italy, the US, UK, and France were recruited by a physician association (ITACA) and patient advocacy groups (HAEA, HAE UK, AMSAO), respectively, to complete an online survey
- Patients had to have treated an attack within the 3 months prior to the survey with an approved on-demand therapy
- The survey was self-reported, and took respondents approximately 20 minutes to complete
- Respondents rated their anxiety using an 11-point GA-NRS ranging from 0 "not anxious" to 10 "extremely anxious" to answer the question "How much anxiety did you feel about treating this HAE attack with on-demand treatment?"
- Attack severity was reported on a 4-point Patient Global Impression of Severity (PGI-S) scale ranging from 1 "Mild" to 4 "Very Severe"

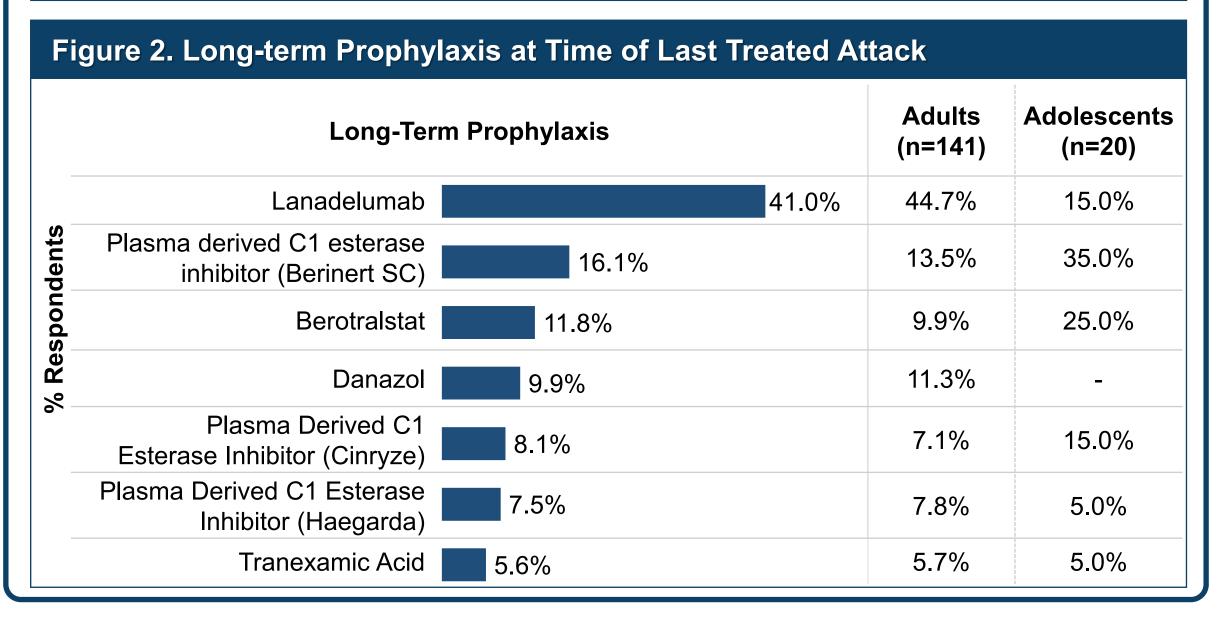
Results

Table 1. Patient Demographics and Clinical Characteristics On-Demand On-Demand Total Adolescents Characteristic (n=31)(N=284)(n=253) (n=179)Current age; years mean 41.0 (16.4) 14.3 (1.6) 33.4 (15.6) 45.4 (15.3) 44.3 (14.3) (SD) Age of diagnosis; years 20.3 (13.1) 17.6 (13.2) 19 (13.3) 6.3 (3.6) 13 (12.1) mean (SD) Gender, n (%) 19 (61.3%) 93 (32.7%) Male 190 (66.9%) 70 (66.7%) 120 (67.0%) 178 (70.4%) 12 (38.7%) Female Country, n (%) 14 (45.2%) 55 (30.7%) 87 (34.4%) Italy 101 (33.1%) 46 (43.8%) 63 (35.2%) **United States** 14 (45.2%) **United Kingdom** 48 (16.9%) 25 (23.8%) 23 (12.8%) 46 (18.2%) 2 (6.5%) 3 (2.9%) 38 (21.2%) 1 (3.2%) 41 (14.4%) 40 (15.8%) France HAE Type, n (%) 258 (90.8%) 161 (89.9%) 27 (87.1%) 97 (92.4%) 231 (91.3%) Type I 8 (7.6%) 18 (10.1%) 26 (9.2%) 22 (8.7%) 4 (13.9%) Type II Days since last attack, 20.7 (19.5) 16.6 (15.8) 23.2 (21.1) 20.6 (19.1) 21.9 (23.4) mean (SD)

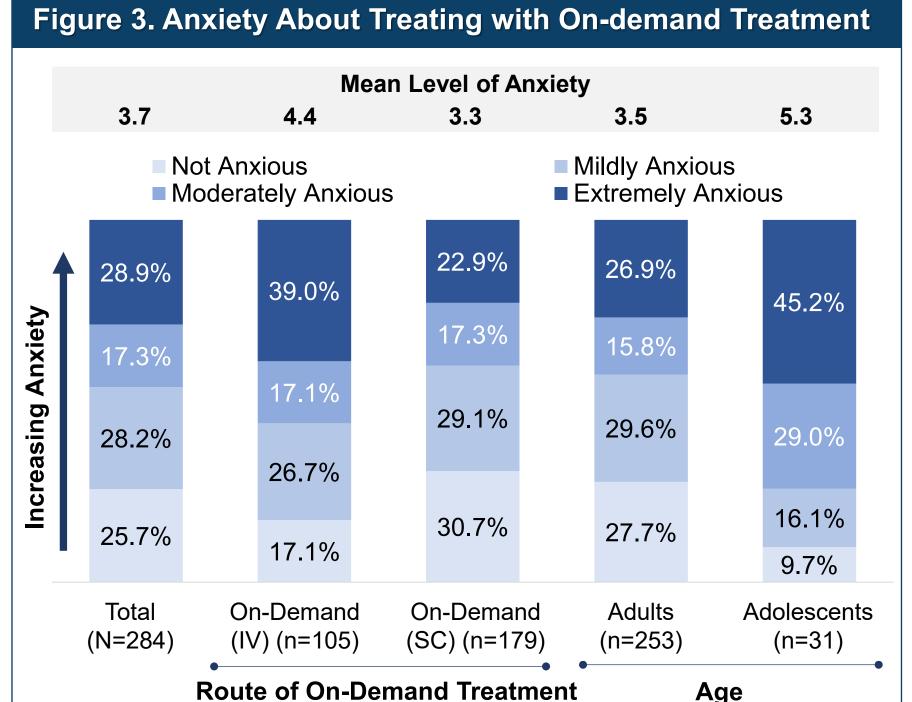
- This analysis included 284 respondents (253 adults [≥18yrs] and 31 adolescents [range 12-17yrs old]) from Italy (n=101), US (n=94), UK (n=48), and France (n=41)
- 57% were receiving long-term prophylaxis

Figure 1. On-Demand Therapy Used for Last Treated Attack					
_	On-Demand Therapy			Adults (n=253)	Adolescents (n=31)
	Icatibant (Firazyr and Generic)		62.3%	68.4%	12.9%
lents	Plasma Derived C1 Esterase Inhibitor (Berinert)	27.1%		22.5%	64.5%
Respondents	Recombinant C1 Esterase Inhibitor (Ruconest)	6.7%		4.7%	22.6%
% Re	Plasma Derived C1 Esterase Inhibitor (Cinryze)	3.2%		3.3%	-
	Ecallantide	0.7%		0.8%	-

87% of adolescents and 31% of adults used an IV on-demand treatment to treat their last attack



Results

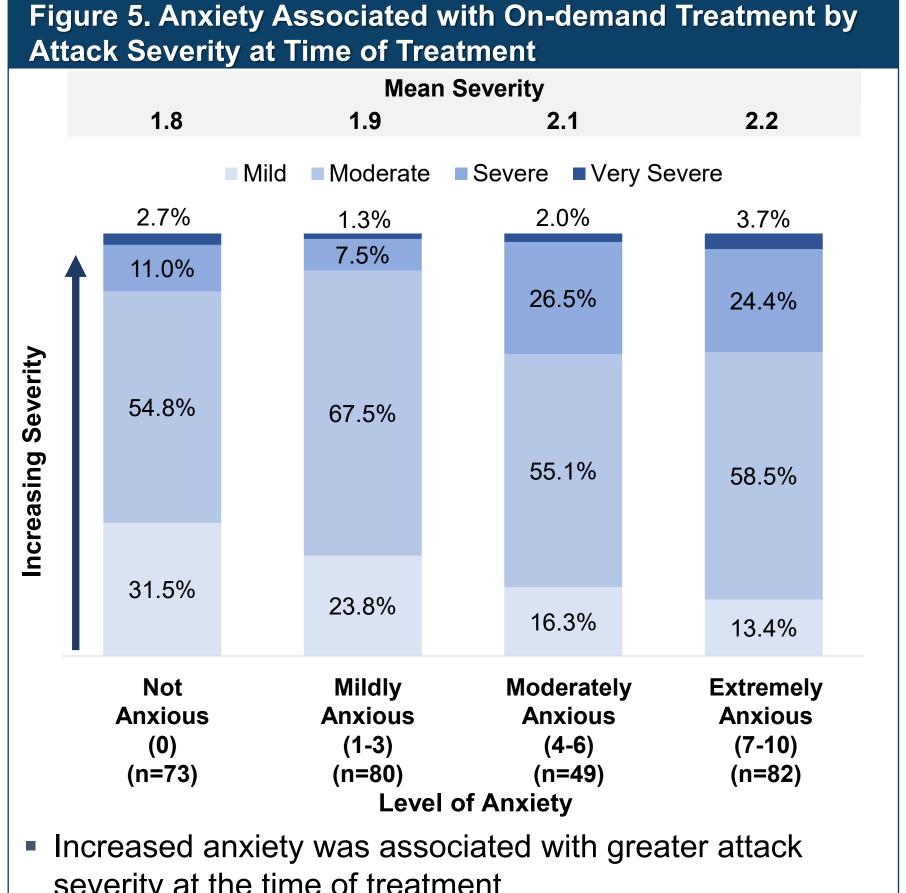


- During the last treated attack, 29% of respondents felt extremely anxious (anxiety 7-10), 17% moderately anxious (anxiety 4-6), and 28% mildly anxious (anxiety 1-3)
- The mean anxiety rating was 3.5 for adults vs 5.3 for adolescents
- For respondents receiving IV treatment, the mean anxiety rating was 4.4, with 39% reporting extreme anxiety

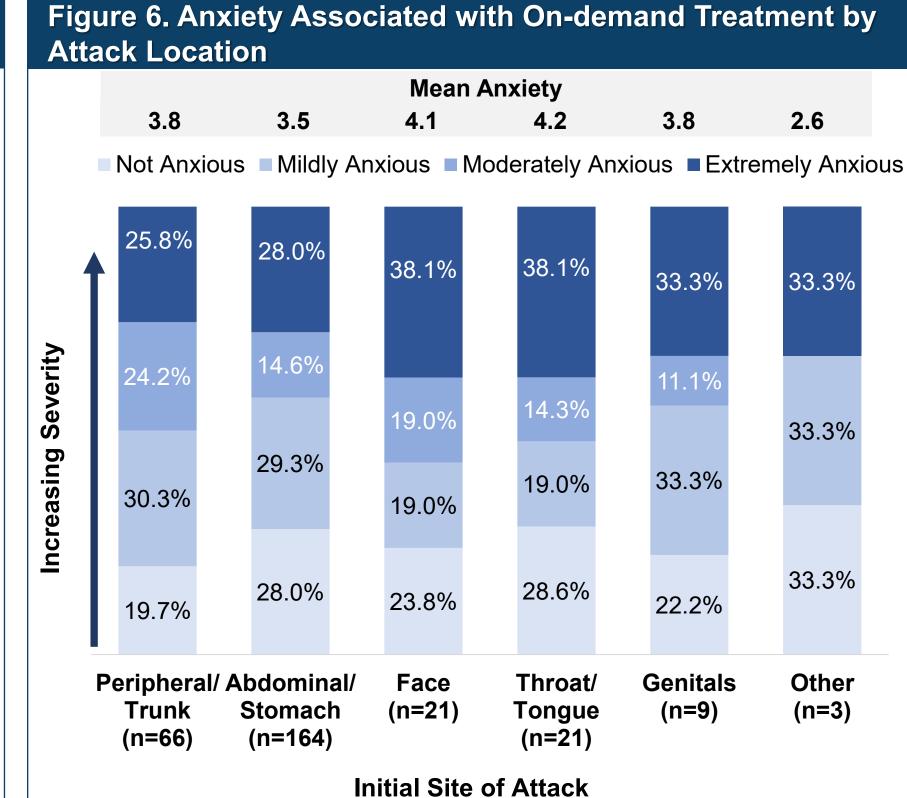
Figure 4. Time to On-demand Treatment and Anxiety **Mean Level of Anxiety** 3.0 3.6 4.3 3.9 ■ Not Anxious (0) ■ Mildly Anxious (1-3) ■ Extremely Anxious (7-10) ■ Moderately Anxious (4-6) 23.1% 24.1% 29.8% 30.8% 41.7% 30.8% 32.8% 26.6% 29.2% 23.1% 23.1% 20.8% ≥8 hours ≥2 to <5 Hours ≥5 to <8 hours (n=39)(n=58)(n=124)(n=24)(n=39)

Time to Treatment

- 16% of respondents treated in <1 hour
- Increased anxiety was associated with increased time to treatment



severity at the time of treatment



 Moderate to extreme anxiety occurred for all attack locations, but was highest in the face or throat/tongue

Figure 7. Reasons for Anxiety Associated with On-demand Treatment <u>Top 5</u> **On-Demand On-Demand Adults** Adolescents **Reasons for Anxiety** (n=183)(n=28)(n=87)(n=124)Desire not to 'waste' an on-demand treatment if the attack was less 44% 39% 7% severe than I thought Uncertainty about how long the treatment would take to begin 24% 36% 32% 21% Running out of on-demand treatment if I needed it later 14% 33% 28% 7% Uncertainty about whether the treatment would work 24% 39% 21% Worry about a rebound attack after the first treatment 26% 14% 21% Anticipating burning or pain with the injection 16% 25% Uncertainty if the attack would become severe enough to treat 16% 16% 25% 36% Finding the vein to start the intravenous infusion 33% 11% 1% The cost of the on-demand treatment 15% 14% 8% The need to use a second dose for the same attack 7% 15% 12% 11% Anticipating side effects from the injection 9% 18% Finding someone to help me administer the treatment 10% 11% 4% 7% 14% I am afraid of needles Finding a private area to administer the treatment 8% 7% 4% 6% 2% 7% The process of preparing my treatment

Finding a vein to start infusion was the most common administration-related reason for anxiety among those receiving IV on-demand treatment (33%) and anticipating burning or pain with injection was the most common administration-related reason for respondents using SC treatment (23%)

Conclusions

- Nearly one third of survey respondents experienced moderate to extreme anxiety due to anticipated use of injectable on-demand treatment, particularly adolescents and those receiving IV therapy
- Greater levels of anxiety were associated with longer on-demand treatment delays and attack severity
- Among the reasons related to treatment administration, finding a vein to start infusion and burning or pain with injection were the most common causes of anxiety
- An oral on-demand therapy could reduce the treatment administration-related causes of anxiety associated with currently approved on-demand therapies

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