

Tryptase testing in suspected systemic allergic reaction

Insights into the connection between transiently elevated tryptase levels and mast cell activation indicating possible anaphylaxis

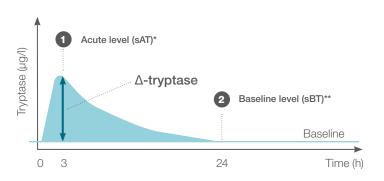
Anaphylaxis is a systemic hypersensitivity reaction usually involving two or more organs including skin/mucus membranes, airways, cardiovascular, and/or gastrointestinal systems. The World Allergy Organization (WAO) provides diagnostic criteria based on clinical parameters.^{1,2} Because anaphylaxis can potentially be life threatening, paired acute and baseline tryptase levels aid in differential diagnosis (including mastocytosis or other mast cell disorders), patient management and follow up care. Tryptase is a useful biomarker as an aid in investigating systemic allergic reactions^{3,4} as it has been shown to be released into the circulatory system during anaphylaxis.^{4,5}

When measuring tryptase levels - timing is important

Leading global allergy authorities such as WAO, AAAAI/ ACAAI and EAACI recommend taking two serum tryptase measurements³⁻⁸ for comparison:

- Acute level (sAT)*: as soon as possible after the clinical reaction onset (+15 minutes up to 3 hours).^{3,4,8,9}
- **Baseline level (sBT)**:** 24-48 hours after complete resolution of all clinical signs and symptoms.^{3,4,8,9}

If the change (Δ) in tryptase levels (sAT - sBT) is \geq 20% of the individual's sBT + 2 µg/l then mast cell activation is indicated.^{2,3,7,8,10} Note that seemingly normal serum tryptase levels do not preclude that an anaphylactic reaction has occurred,^{2-5,7,9,11} so performing the calculation can help identify possible mast cell activation that might otherwise be missed.^{8,11}

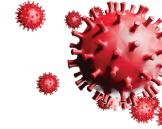


If mast cell activation is indicated, the WAO and EAACI recommend referral to a specialist (e.g., allergist, immunologist) for follow-up, along with other measures such as counseling, before patient discharge from the acute care setting.^{2,5,6,8,9,12}



Measure tryptase levels twice

when systemic allergic reaction is suspected



Systemic allergic reaction & COVID-19 vaccines

Due to the global COVID-19 pandemic and rollout of vaccine initiatives, it is important to have increased awareness that, to date, the incidence rate of anaphylaxis associated with first doses of three major COVID-19 vaccines exceeds the estimated incidence rate of anaphylaxis associated with other vaccines (2 to 11.1 per million¹³⁻¹⁶ vs approximately 1 per million^{12,16-18}).

Guidelines for tryptase testing for suspected COVID-19 vaccine-associated anaphylaxis

Global allergy organizations such as WAO and EAACI-ARIA issued statements and/or guidelines^{12,16-19} on the diagnosis, management, and prevention of severe allergic reactions to COVID-19 vaccines. As patient history and symptomology is often ambiguous, tryptase testing is critically important for accurate diagnosis and to minimize future risk of severe allergic reactions in vaccinated individuals.¹⁷ As is the case with other anaphylactic events, the same organizations recommend affected individuals follow-up with a specialist (e.g., allergist, immunologist) to determine if a second dose of COVID-19 vaccine should be given.^{12,16,19}

References

- 1. Beck SC *et al.* Biomarkers in Human Anaphylaxis: A Critical Appraisal of Current Evidence and Perspectives, *Frontiers in Immunology* 2019; 10:494
- Simons FE et al. International consensus on (ICON) anaphylaxis. World Allergy Organ J. 2014 May;30;7(1):9
- Valent *et al.* Definitions, criteria, and global classification of mast cell disorders with special reference to mast cell activation syndromes: a consensus proposal. *Int Arch Allergy Immunol.* 2012;157(3):215–25.
- Schwartz LB. Diagnostic value of tryptase in anaphylaxis and mastocytosis. *Immunol Allergy Clin North Am.* 2006 Aug;26(3):451–63.
- 5. Lieberman *et al.* The diagnosis and management of anaphylaxis practice parameter: 2010 update. *J Allergy Clin Immunol.* 2010 Sep;126(3):477-80.
- Muraro *et al.* Anaphylaxis: guidelines from the European Academy of Allergy and Clinical Immunology. *Allergy*. 2014 Aug;69(8):1026-45.
- Cardona *et al.* World allergy organization anaphylaxis guidance 2020. World Allergy Organ J. 2020 Oct 30;13.
- Vitte *et al.* Use and Interpretation of Acute and Baseline Tryptase in Perioperative Hypersensitivity and Anaphylaxis. *J Allergy Clin Immunol.* 2021 Mar 18;S2213-2198(21)00315-9. DOI: 10.1016/j.jaip.2021.03.011.
- Schwartz LB, Yunginger JW, Miller JS *et al.* The time course of appearance and disappearance of human mast cell tryptase in the circulation after anaphylaxis. *J Clin Invest.* 1989 May;83(5):1551–5.
- Simons FE et al. World Allergy Organization anaphylaxis guidelines: Summary. J Allergy Clin Immunol. 2011 Mar;127(3):587-93.
- 11. Simons FE et al. 2015 update of the evidence base: World Allergy Organization anaphylaxis guidelines. *World Allergy Organ J.* 2015 Oct 28;8(1):32.
- 12. Sokolowska *et al.* EAACI statement on the diagnosis, management, and prevention of severe allergic reactions to COVID-19 vaccines. *Allergy.* 2021 Jan 16.

- Allergic Reactions Including Anaphylaxis After Receipt of the First Dose of Pfizer-BioNTech COVID-19 Vaccine — United States, December 14–23, 2020. MMWR Morb Mortal Wkly Rep 2021; 70:46–51. DOI: http://dx.doi.org/10.15585/mmwr.mm7002e1
- Allergic Reactions Including Anaphylaxis After Receipt of the First Dose of Moderna COVID-19 Vaccine - United States, December 21, 2020–January 10, 2021. MMWR Morb Mortal Wkly Rep 2021; 70:125–129. DOI: http://dx.doi.org/10.15585/mmwr. mm7004e1
- COVID-19 vaccine safety update for Vaxzevria, AstraZeneca AB. :(2021, March 29). European Medicines Agency. https://www.ema.europa.eu/documents/ covid-19-vaccine-safety-update/covid-19-vaccine-safety-update-vaxzevria-previouslycovid-19-vaccine-astrazeneca-29-march-2021_en.pdf
- American College of Allergy, Asthma, and Immunology (ACAAI). (2021, March 11). ACAAI Updates to Guidance on Risk of Allergic Reactions to COVID-19 Vaccines. [Press Release]. https://acaai.org/news/acaai-updates-guidance-risk-allergic-reactions-covid-19-vaccines
- Turner *et al.* COVID-19 vaccine-associated anaphylaxis: A statement of the World Allergy Organization Anaphylaxis Committee. *World Allergy Organ J.* 2021 Feb;14(2):100517.
- Klimek *et al.* ARIA-EAACI statement on severe allergic reactions to COVID-19 vaccines

 an EAACI-ARIA Position Paper. *Allergy.* 2020 Dec 30.
- US Centers for Disease Control and Prevention (CDC), referred to by AAAAI. (2021, April 8). Lab Tests to Collect Shortly After Severe Allergic Reaction/ Anaphylaxis Following COVID-19 Vaccination https://www.cdc.gov/ vaccines/covid-19/clinical-considerations/testing-after-allergic-reaction. html?CDC_AA_refVal=https%3A%2F%2Fwuww.cdc.gov%2Fvaccines%2Fcovid-19%2Fclinical-considerations%2Fanaphylaxis-management.html

Find more resources at thermofisher.com/measuretryptasetwice

The cleared intended use of ImmunoCAP Tryptase differs in the United States.

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