

## Yuki Nakamura

## Personal Information

Date of Birth: 11/11/1982

Nationality: Japanese



## Work Experience

2010- Present Assistant Professor, Department of Immunology, Interdisciplinary Graduate School of Medicine and Engineering, University of Yamanashi

2009- 2010 Research Scientist, R&amp;D Placement Business Division, World Intec Co., Ltd.

## Education

2014 Ph.D. in Medical Science, University of Yamanashi

2009 M.S. in Medical Science, University of Yamanashi

2007 B.S. in Engineering, University of Yamanashi

## Awards

2011 7th Annual Meeting of the Japanese Society of Allergology Scientific Congress Award

2011 21st Abbott Japan Allergy Academic Award

## Research Interest

Circadian rhythm, Mast cell

## Publication

1. Nakamura Y et al: Circadian regulation of allergic reaction by the mast cell clock in mice. *J Allergy Clin Immun* 133(2):568-575.e12, 2014.
2. Nakamura Y et al: Circadian clock gene *Period2* regulates a time-of-day-dependent variation in cutaneous anaphylactic reaction. *J Allergy Clin Immun* 127(4):1038-1045, 2011.
3. Nakamura Y et al: The latent form of transforming growth factor- $\beta$  administered orally is activated by gastric acid in mice. *J Nutr* 139(8):1463-1468, 2009.
4. Nakamura Y et al: House dust mite allergen Der f 1 can induce the activation of latent TGF- $\beta$  via its protease activity. *FEBS Letters* 583(12):2088-2092, 2009.
5. Nakamura Y et al: Cigarette smoke extract induces thymic stromal lymphopoietin expression, leading to TH2-type immune responses and airway inflammation. *J Allergy Clin Immun* 122(6):1208-1214, 2008.

## Circadian regulation of allergic reaction by the mast cell clock in mice

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**Background:** It remains elusive how allergic symptoms exhibit prominent 24-hour variations. Recently, we have shown that the circadian clocks drive the daily rhythms in IgE/mast cell-mediated allergic reactions. However, the precise mechanisms, particularly the specific roles of the mast cell-intrinsic clockwork in the temporal regulation, remain unclear.

**Objective:** We determined whether the mast cell clockwork contributes to the temporal regulation of IgE/mast cell-mediated allergic reactions.

**Methods:** The kinetics of a time-of-day-dependent variation in passive cutaneous anaphylactic (PCA) reaction were compared between mast cell-deficient mice reconstituted with bone marrow-derived mast cells (BMMCs) generated from mice with wild-type allele and a non-functional mutation of a key clock gene *Clock*. In addition, we examined the temporal responses of wild-type and *Clock*-mutated BMMCs to IgE stimulation in vitro.

**Results:** The *Clock*-mutation in mast cells resulted in the absence of temporal variations in IgE-mediated degranulation in mast cells both in vivo and in vitro in association with the loss of temporal regulation of Fc $\epsilon$ R1 signaling in mast cells.

**Conclusion:** The mast cell-intrinsic clockwork primarily contributes to the temporal regulation of IgE/mast cell-mediated allergic reaction. The results reveal a novel regulatory mechanism for IgE-mediated mast cell response that may underlie the circadian pathophysiology in allergic diseases.