

# Mechanisms of Allergen-specific immunotherapy

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DAVOS

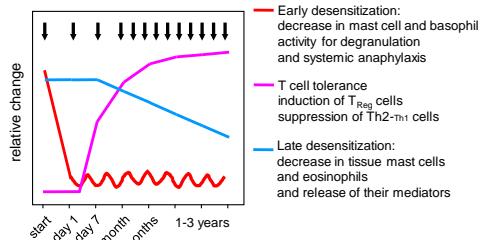
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## mechanisms of SIT

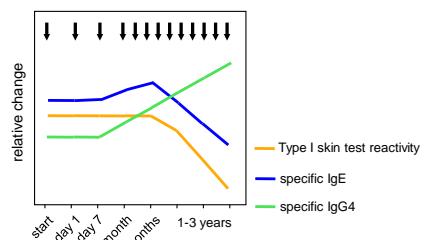


1. Basophil, mast cell desensitisation, early events related to APC (DC, B cell), effect of adjuvants
2. Regulation of T cells and related phenomena (Treg cells and allergen tolerance)
3. Regulation of B cells, antibodies and related phenomena (Breg cells and IgG4 in allergen tolerance)

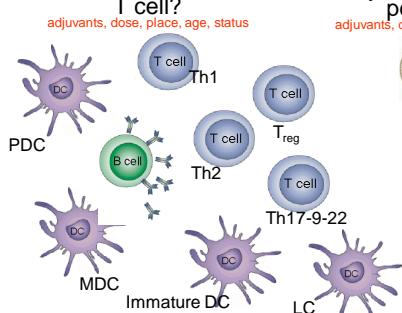
## Mechanisms of allergen-specific immunotherapy



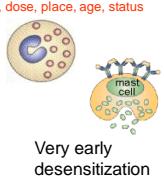
## Mechanisms of allergen-specific immunotherapy



Q1: which one will be the first and preferential APC to present the allergen which T cell?



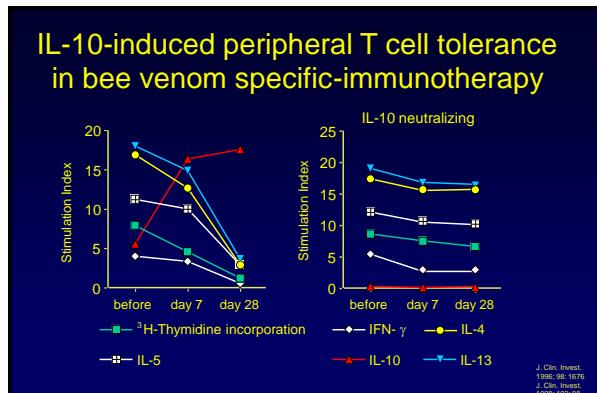
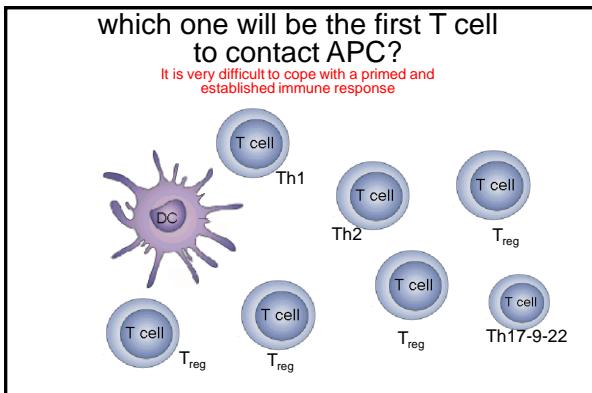
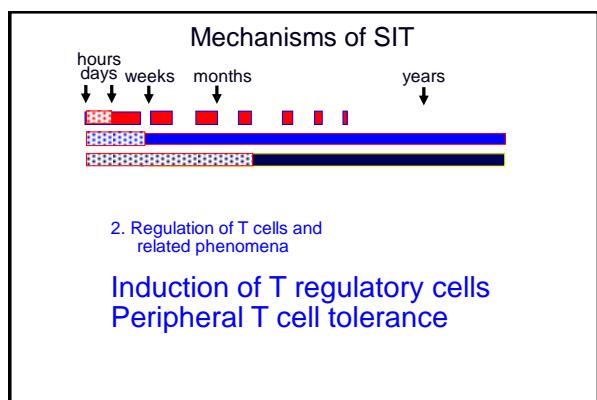
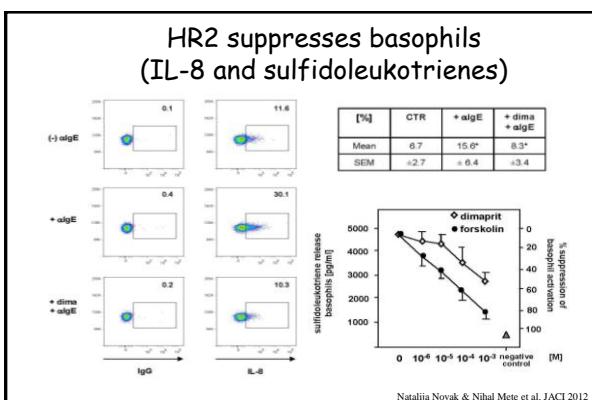
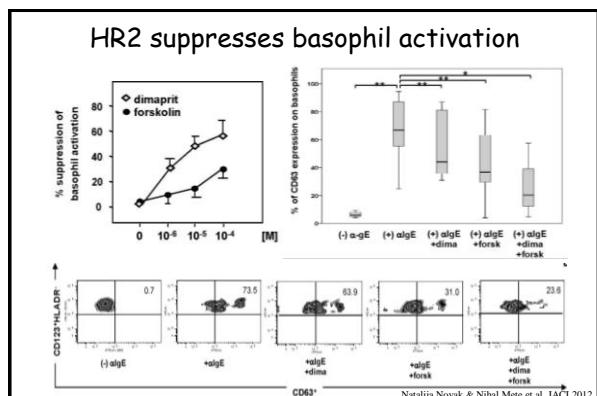
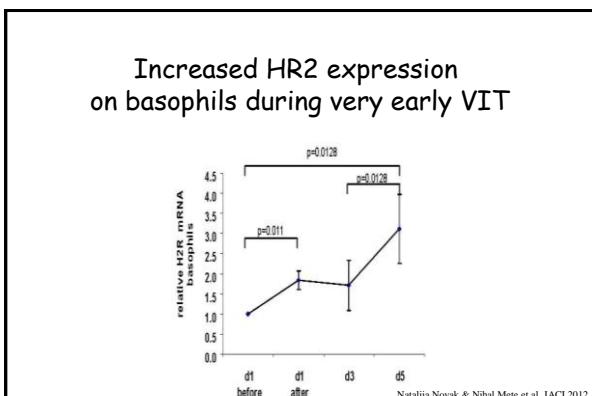
Q2: how will the system avoid anaphylaxis as early and efficient as possible?

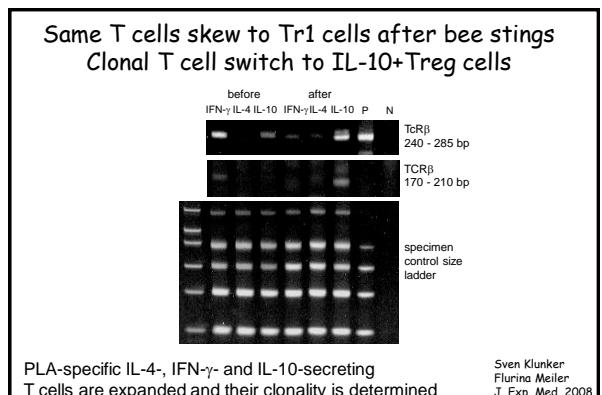
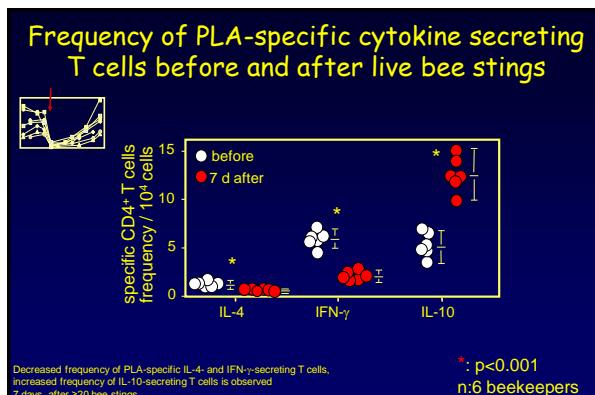
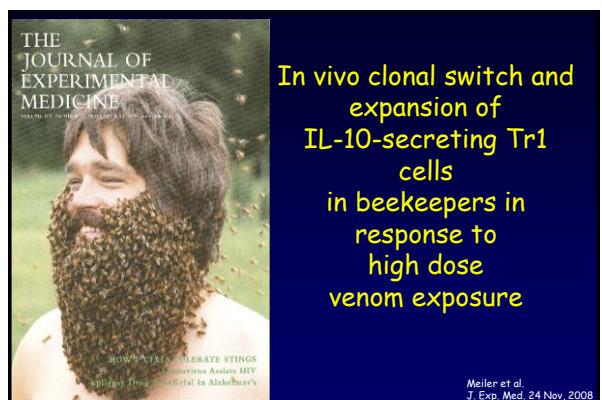
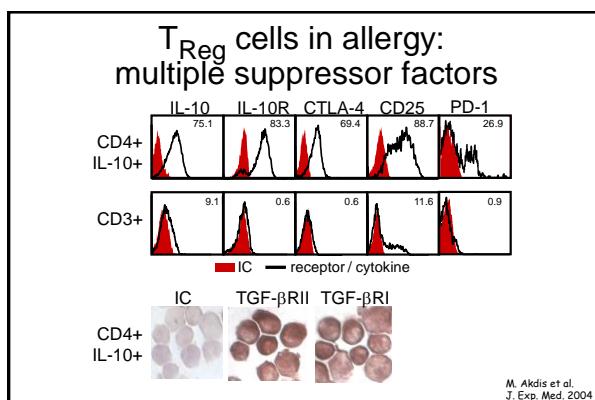
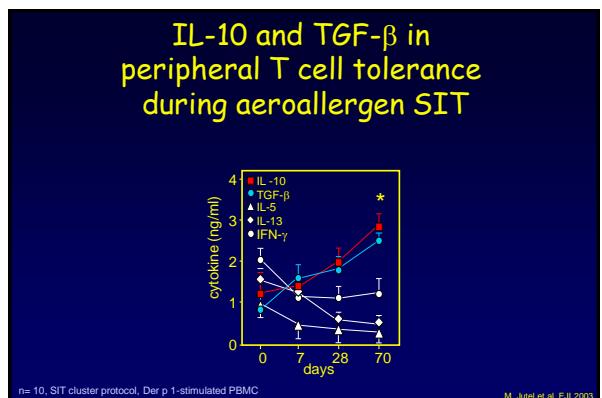
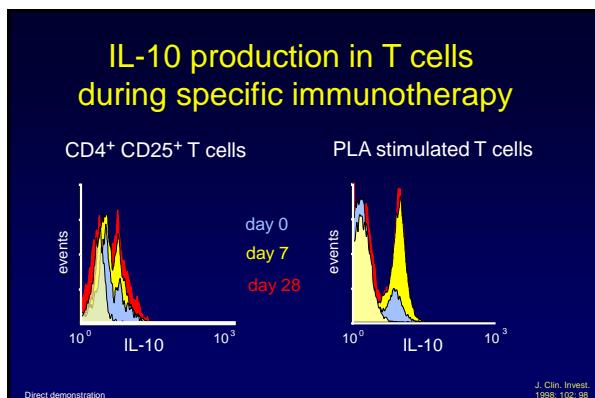


## mechanisms of SIT (very early events)

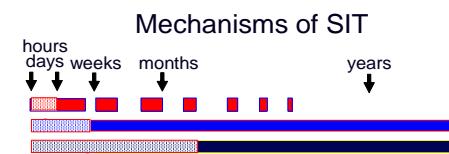


1. Basophil, mast cell desensitisation,





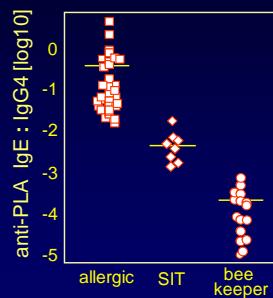
# Regulation of B cells, antibodies and related phenomena



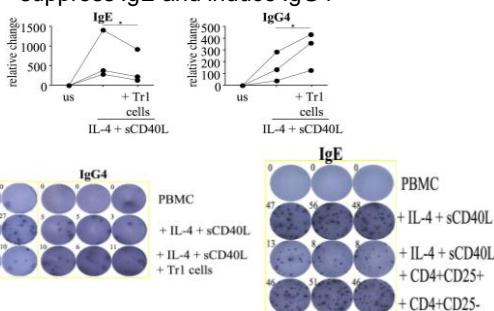
3. Regulation of B cells, antibodies and related phenomena



PLA-specific IgE:IgG4 antibody ratios



IL-10, Tr1 and FoxP3+Treg cells suppress IgE and induce IgG4

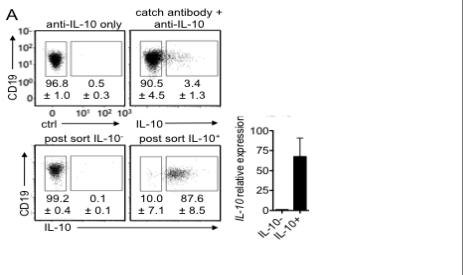


Meiler et al. Allergy 2008

Human Breg cells in allergen tolerance and allergen-SIT

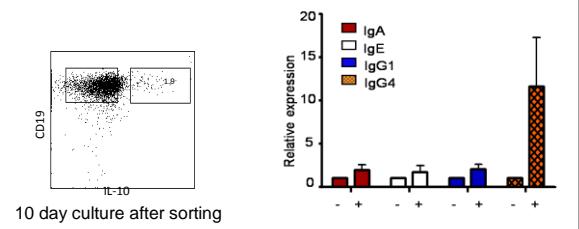
Willems van de Veen et al.  
M. Akdis

## Circulating Human IL-10+ Breg cells



Willem van de Veen et al.  
M. Akdis

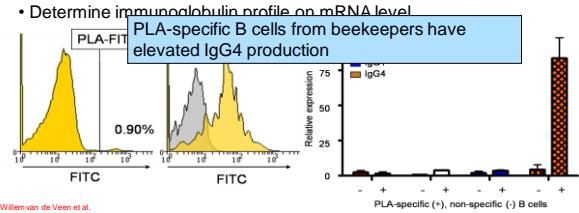
## IgG4 production in IL-10-producing B cells



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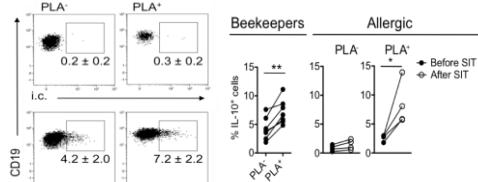
## Allergen-specific B cells produce IgG4 in allergen tolerance

- Isolate CD19+ B cells from beekeepers
- Stain with PLA-FITC
- Sort cells
- Determine immunoglobulin profile on mRNA level



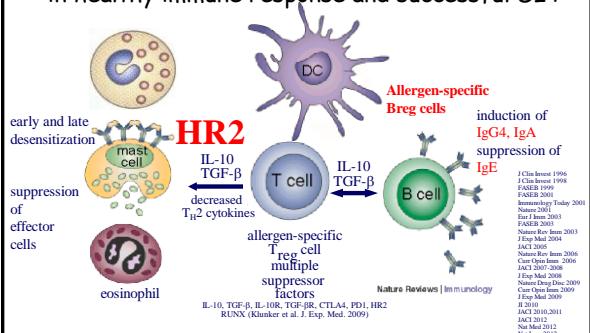
Willem van de Veen et al.  
M. Akdis

## Increased IL-10-producing Breg cells during allergen-SIT



Willem van de Veen et al.  
M. Akdis

## Immune tolerance to allergens in healthy immune response and successful SIT





<ul style="list-style-type: none"> <li>• Oscar Palomares</li> <li>• Nihal Mete</li> <li>• Natalija Novak</li> <li>• Judith Zumkehr</li> <li>• Flurina Meiler</li> <li>• Sven Klunker</li> <li>• Alison Taylor</li> <li>• Johan Verhagen</li> <li>• Andrea Joss</li> <li>• Maya Zimmermann</li> <li>• Christian Manhart</li> <li>• Tomasz Basinski</li> <li>• Tunc Akkoc</li> <li>• Axel Trautmann</li> <li>• Mübellel Akdis</li> <li>• Reto Cramer</li> <li>• Liam O'Mahony</li> <li>• Roger Lauener, Davos</li> </ul>	<p><b>Immune tolerance</b></p> <ul style="list-style-type: none"> <li>• U. Müller, Bern</li> <li>• M. Jutel, Wroclaw, SIAF</li> <li>• T. Kündig, Zürich</li> <li>• G. Senni, Zürich</li> <li>• B. Wüthrich, Zurich</li> <li>• Natalija Novak, Bonn</li> <li>• R. Valenta, Vienna</li> <li>• M. van Hage, Stockholm</li> <li>• G. Gavvelin, Stockholm</li> <li>• M. Colonna, St Louis</li> <li>• E. Flory, Langen</li> <li>• S. Viehts, Langen</li> <li>• H. Fiebig, Reinbek</li> <li>• O. Cromwell, Reinbek</li> <li>• E. Hamelmann, Berlin</li> <li>• M. Larché, London</li> <li>• Ch. Heusser, Basel</li> <li>• B. Ballmer-Weber, Zurich</li> </ul>	<p><b>Effector Mechanisms</b></p> <ul style="list-style-type: none"> <li>• R. Disch</li> <li>• B. Wüthrich</li> <li>• P. Schmid-Grendelmeier</li> <li>• W. Kneist</li> <li>• M. Schilz</li> <li>• D. Kleeman</li> <li>• W. Deglmann</li> <li>• H. Behrendt.</li> <li>• C. Traidl-Hoffmann,</li> <li>• G. Menz</li> </ul>
	<p><b>Immune regulation by histamine</b></p> <ul style="list-style-type: none"> <li>• T. Watanabe,</li> <li>• R. Koga, Fukuoka</li> <li>• M. Jutel, Wroclaw</li> </ul>	

