

# The role of FRs in indoor dust in potentiating/facilitating allergic responses to inhaled allergens

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#### Aim



- » A tremendous increase in both indoor exposure to FRs and prevalence of allergy over the last decades
- » A major exposure route to FRs via inhalation of house dust which also contains allergens, like HDM

THAT'S WHY:

"Investigating possible roles and effects of FRs on the immune response to inhaled allergens in asthma"

# 6/27/2012

### Allergic asthma

#### INM Institute for Environmental Studies

#### » Caused by:

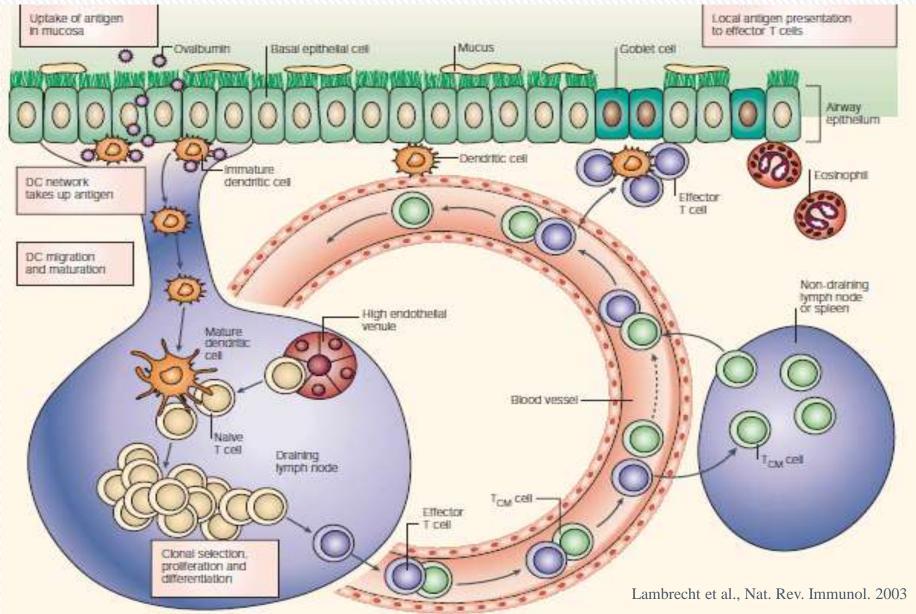


#### Characterized by:





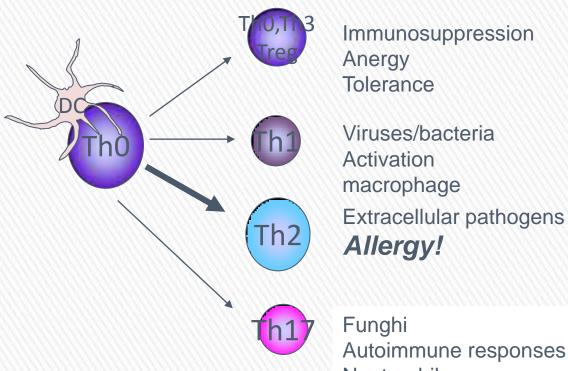
## The immune response to inhaled allergen



# T cell stimulation by DCs



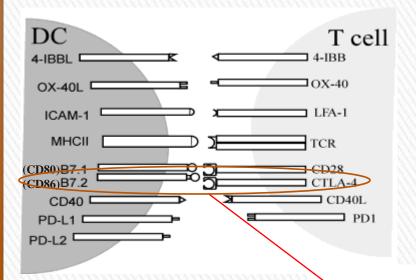
> Activated DCs can determine the type of the immune response by skewing naïve T cells towards:



Autoimmune responses Neutrophil recruitment

#### How?



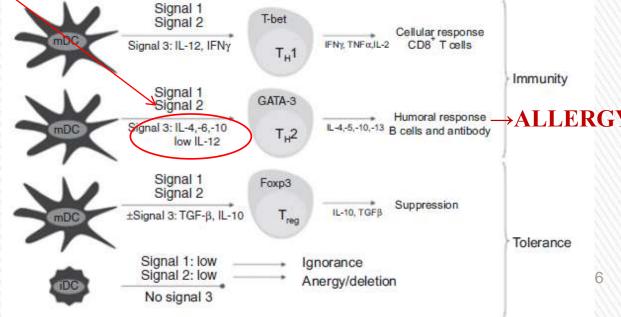


**Signal 0:** Interaction of antigen with DC

**Signal 1:** Antigen presentation by ligation of MHCII-antigen with TCR

**Signal 2:** Activation of T cells by costimulatory molecules

**Signal 3:**Differentiation of T cells by cytokines







### Preliminary experiments (in vitro)

Setup:



**BONE MARROW ISOLATION + 10 DAYS GMCSF** 



DENDRITIC CELLS; PULSE WITH ANTIGEN (HDM) EXPOSE TO FLAME RETARDANT (HBCD)



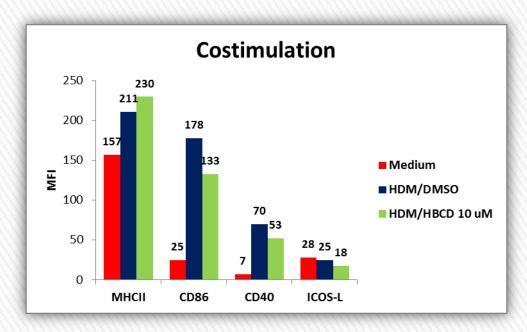
FACS ANALYSIS FOR DETECTING COSTIMULATORY MOLECULES + ELISA OF SUPERNATANTS FOR CYTOKINE PRODUCTION

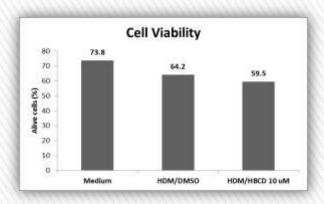


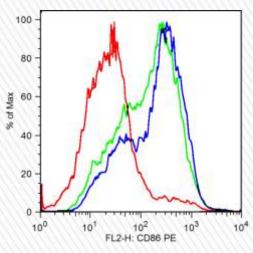


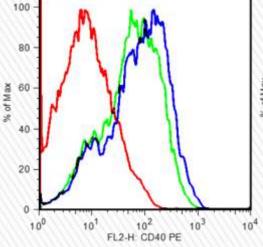


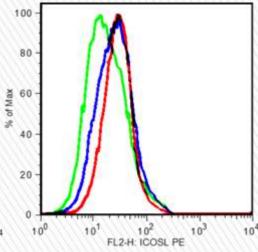
#### **Results (FACS)**









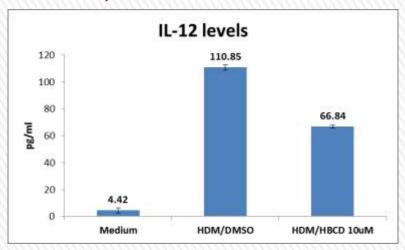


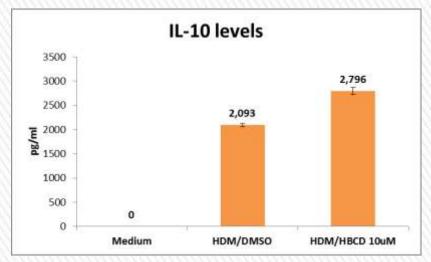


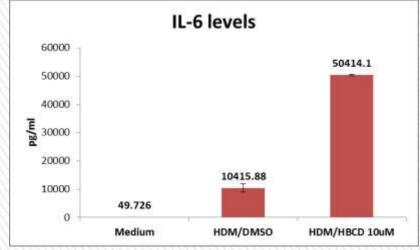




# **Results (ELISA)**











#### **Conclusion**

Effects of HBCD/ HDM Exposure			
	Molecule	Level	Sensitization
Costimulatory	CD86		Anti-allergy
	CD40		Pro-allergy
	ICOS-L		Anti/pro allergy
Cytokine	IL-12		Pro-allergy
	IL-6	<u> </u>	Pro-allergy
	IL-10	<u></u>	Pro-allergy

# The NET EFFECT on immune response: INDEFINITE!



## **Future plans**





#### FRs monitored in INFLAME WPs 1 and 2

#### Group 1

PBDEs 28, 47, 99, 100, 153, 154, 183, and 209

Tetrabromobisphenol-A (TBBP-A)

Hexabromocyclododecane (HBCD - a, b, and g- diastereomers)

Decabromodiphenylethane (DBDPE)

1,2-bis(2,4,6-tribromophenoxy)ethane (BTBPE)

2-ethylhexyl 2,3,4,5-tetrabromophthalate (TBPH)

2-ethylhexyl 2,3,4,5-tetrabromobenzoate (TBB)

Triphenyl phosphate (TPP)

Tri(2,3-dichloropropyl) Phosphate (TDCPP)

Bor Tri(2-butoxyethyl)phosphate (TBEP)

GM Tri(2-chloroethyl)phosphate (TCEP)

Tri(monochloropropyl)phosphate (TCPP)

#### Group 2

Tetrabromobisphenol A-bis(2,3-dibromopropylether (TBBPA-DBPE)

Syn-dechlorane plus (syn-DP)

Anti-dechlorane plus (anti-DP)

De Tris(2,3-dibromopropyl)isocyanurate (TDBPIC)

FA Pentabromobenzylalcohol (PBBA)

Brominated polystyrene EL

Pentabromotoluene (PBT)

Pentabromethylbenzene (PBEB)

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# THANK YOU FOR LISTENING ©

