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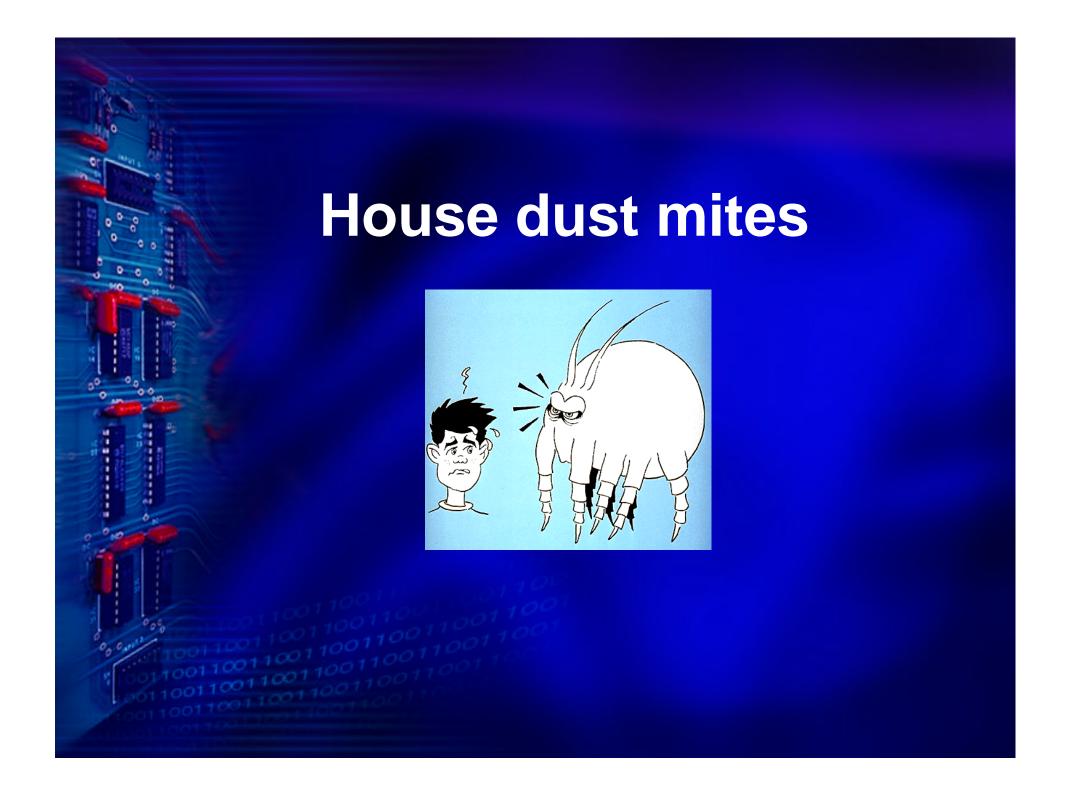
There is now strong evidence that sensitisation and exposure to indoor allergens is a primary cause for asthma

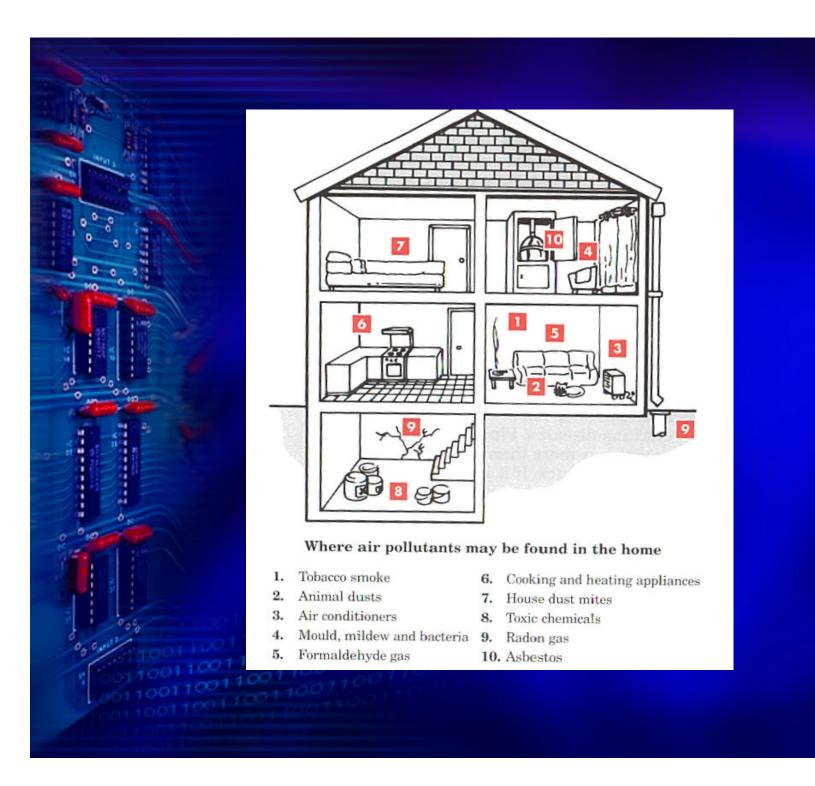
Over the last 4 decades population of the developed world retreated indoors.

Homes have become tighter and more energy efficient ideal for house dust mite growth.

Pet ownership increased

Substandard housing in inner cities associated with cockroach infestation





#### Criteria for causality

In 1965, Hill postulated criteria to be fulfilled in order to demonstrate that an association represents a causal relationship.

In 1992 Sporik applied the criteria of Hill to the association of dust mites and asthma.



Strength: the strength of the association is large

Consistency: repeated observations in different populations have consistent findings

Specificity: a cause leads to a specific effect

Temporality: a cause precedes an effect

Biological gradient: there is a dose-response gradient

**Experiment:** there is experimental evidence

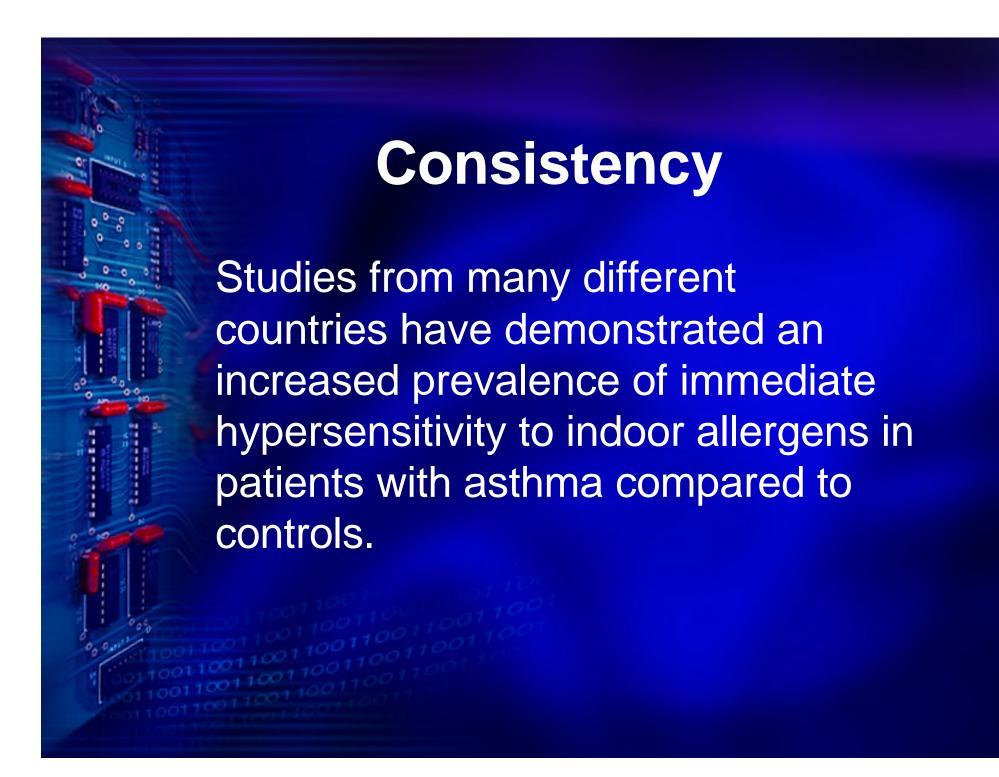
Analogy: there are analogous explanations

Plausibility and coherence: the mechanism is biologically plausible

# The strength of the association is large

Up to 85% of patients with asthma are skin prick sensitive to house dust mites, as compared to 5-30% of the general population.

Importance of sensitisation to allergens of domestic pets - data from Los Alamos, New Mexico, USA (altitude - 2,195m) and Scandinavia.



#### Specificity

Indoor allergens do not appear to be implicated in the aetiology of any other disorder apart from asthma

#### A cause precedes an effect

- Until the early 1970s asthma was almost nonexistent in Papua New Guinea with a prevalence of 0.1
- After the arrival of missionaries with blankets infested with mites prevalence of asthma rose to 7.3.



Exposure and sensitization – already proven for mites

The higher the level of exposure the higher the risk of sensitization.

Kuher J et al. *J Allergy Clin Immunol* 1994;94:44-52 Lau S, et al. *J Allergy Clin Immunol* 1989;84:718-725. Peat et al. *Am J Respir Crit Care Med* 1996;152:144-146.

Exposure and the severity of asthma More difficult to prove...

Peat et al. *Am J Respir Crit Care Med* 1996;152:144-146. Custovic et al. *J Allergy Clin Immunol* 1996;98:64-72.



Reducing the exposure of patients improve their asthma control

Studies in which patients were removed from their homes:

To the mountains with reductions in BHR to histamine — Kerrebijn et al. 1970.

To the hospital — Platts-Mills et al. *The Lancet*,1982.

Studies in which allergen avoidance measures were applied to patients homes

Using mattress encasing — Ehnert B et al. J Allergy Clin Immunol;1993.

## The analogous explanations

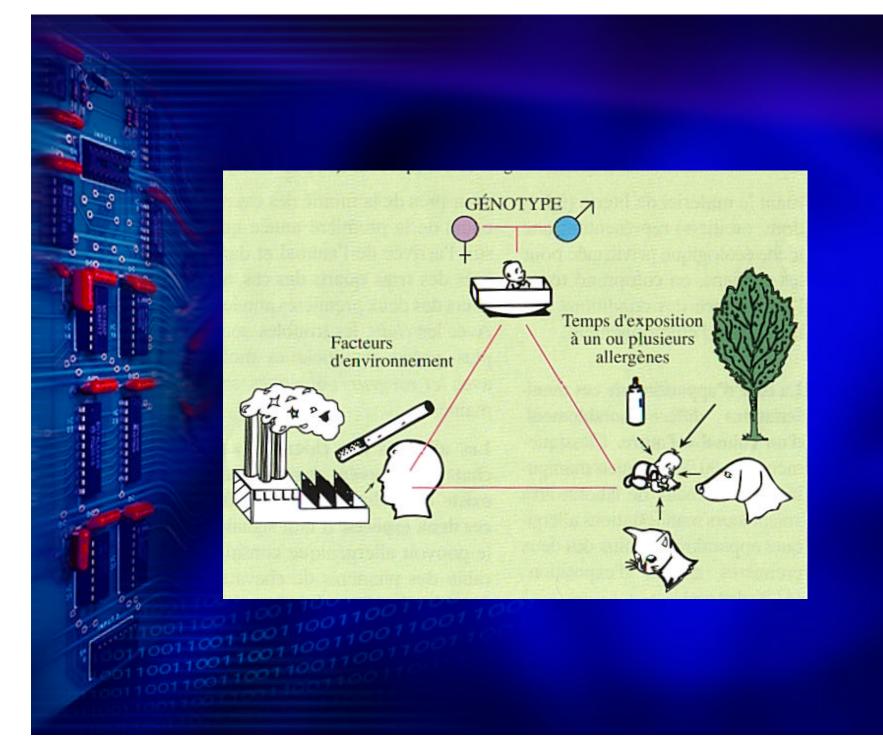
In occupational asthma exposure to allergenic proteins present in the workplace induces asthma in the same way allergenic proteins present in the home do.

The treatment of choice is removal from the workplace resembling allergen avoidance in asthmatics sensitized to indoor allergens.

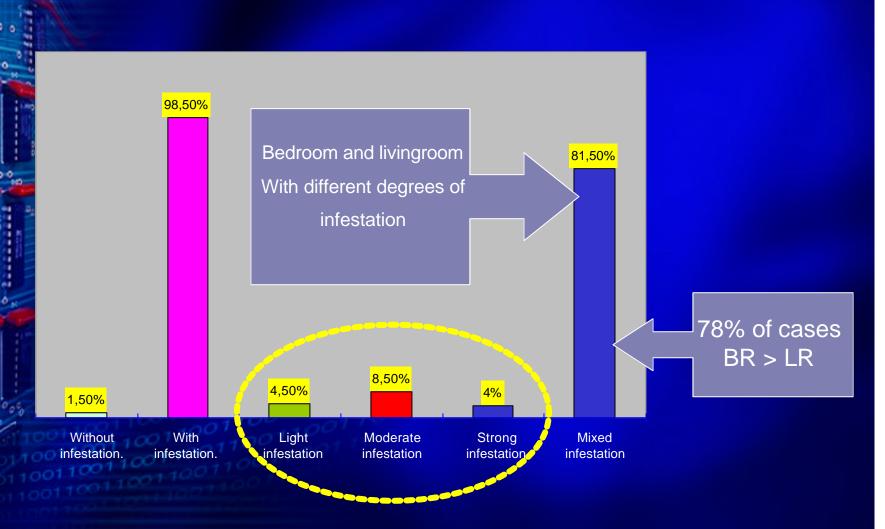
### The mechanism is biologically plausible

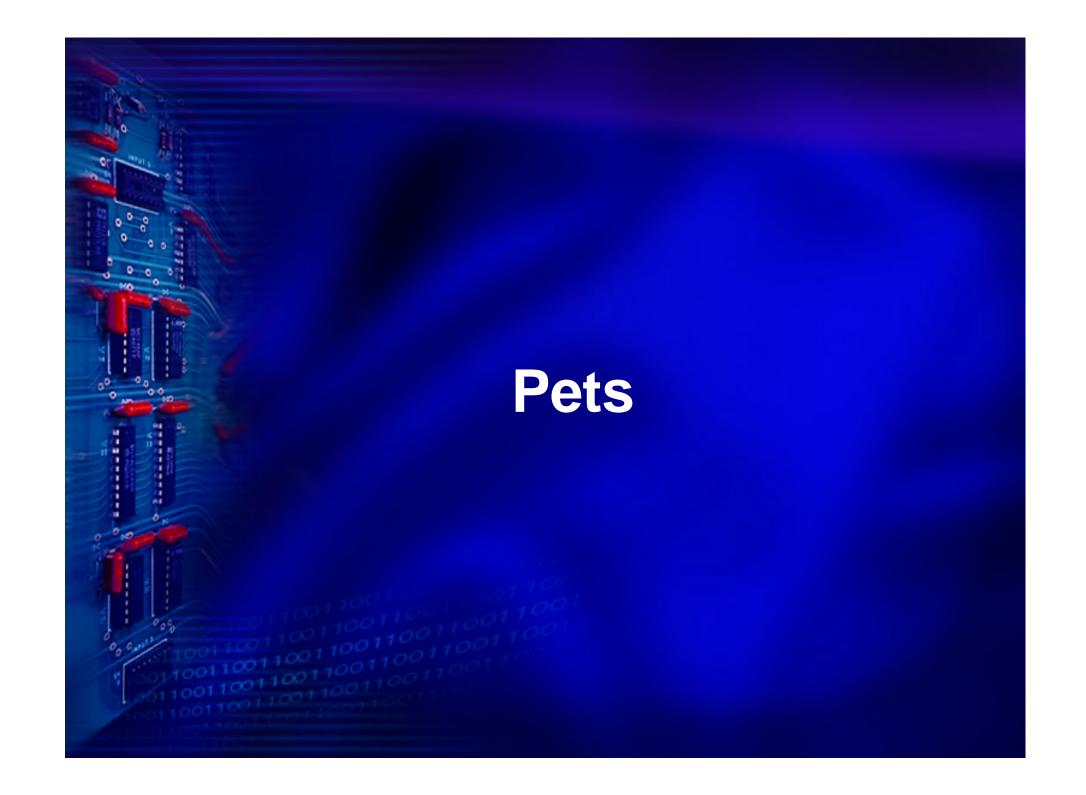
Asthma occurs as a result of a complex interaction between genetic predisposition and the environment.

If predisposed, individuals who are continuously exposed to the allergens they are sensitized, may be at risk of developing airway inflammation and BHR which are the key features of asthma.









|  | Major Cat allergen             | Major Dog Allergen |
|--|--------------------------------|--------------------|
|  | (Fel d I)                      | (Can f 1)          |
| Main   | Anal sacs                      | Dander             |
| sources  | Sebaceous glandes              | Saliva             |
|  | Saliva                         | Skin               |
| Production   | <b>Hormonal control</b>        | Diff. breeds       |
|  | <b>Male &gt; concentration</b> | Same breed         |
|  | High / low producers           |                    |
| <b>Exposition %</b>  | 21.8 - 30.8 %                  | 31 - 50.2 %        |
| Airborne   | 25 % < 2.5                     | ~ 20 % < 5         |
| (microns)  |                                |                    |
|  |                                |                    |
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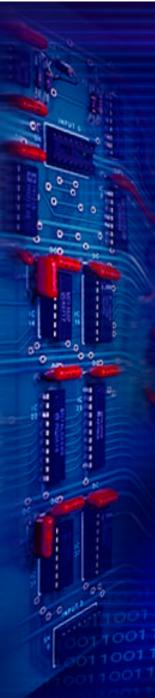
**RISK OF SENSITIZATION.** 

(3 rd International Workshop of Indoor Allergens and Asthma. Cuenca, Spain -July 1995)



The unification of Germany and the impact of environment on the development of allergies in two ethnically similar populations.

von Mutius E, et al. Am J Respir Crit Care Med 1994



### Prevalence of asthma in West and East Germany

The prevalence of doctor diagnosed asthma, hay fever and atopy was higher in West Germany.

Respiratory infections were higher in East Germany.

Day care establishments were attended by 69-71% of East German children aged 1-3 years whereas only 6.9-8.2% of all children in this age group had access to day care in West Germany.

The prevalence of atopic sensitisation decreased significantly with increasing numbers of siblings.

High rates of road traffic impaired lung function and and increased respiratory symptoms in children living in West Germany, but didn't increase the prevalence of rhinitis and asthma.

Several other authors reported similar results.

An East-West European gradient is likely to exist.



These findings suggest that outdoor air pollution is not a strong determinant for the development of asthma and atopy.

It seems unlikely that differences in allergen exposure may explain the difference in atopic sensitisation between East and West Germany.

Factors such as early childhood exposure to viral infections could play a role in the difference of the prevalence of atopic sensitisation between East and West Germany.