

# Rubber, new allergens and preventive measures

© Dr Crépy MN APHP



Marie-Noëlle CREPY  
Paris University Hospital,  
Centre Hôtel-Dieu

1) diagnosis :



# Frequent

- Rubber : one of the main causes of occupational contact dermatitis,
- mostly rubber gloves, especially in health care workers, cleaners and construction workers.

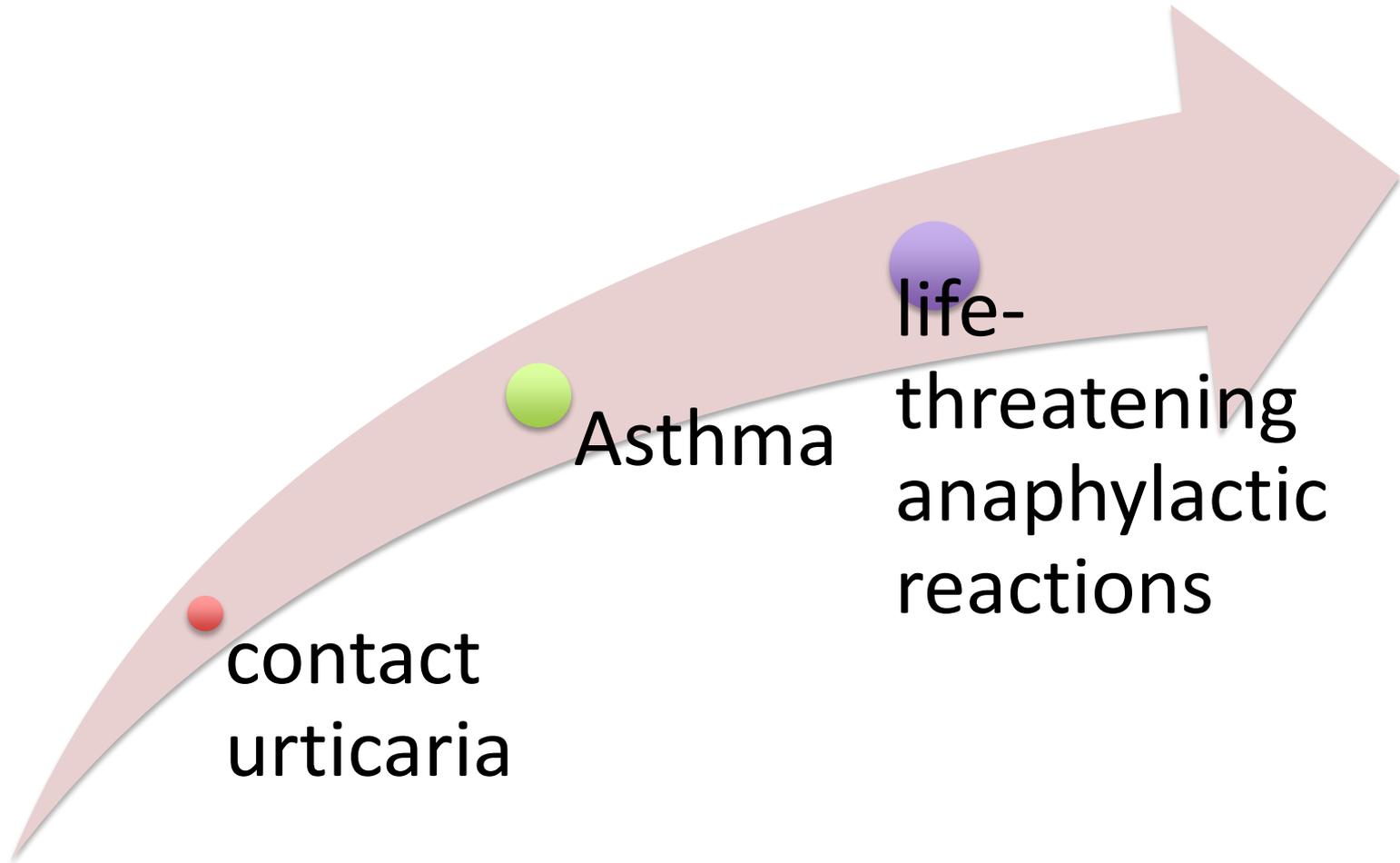
Immediate :  
Type I

Delayed  
Type IV

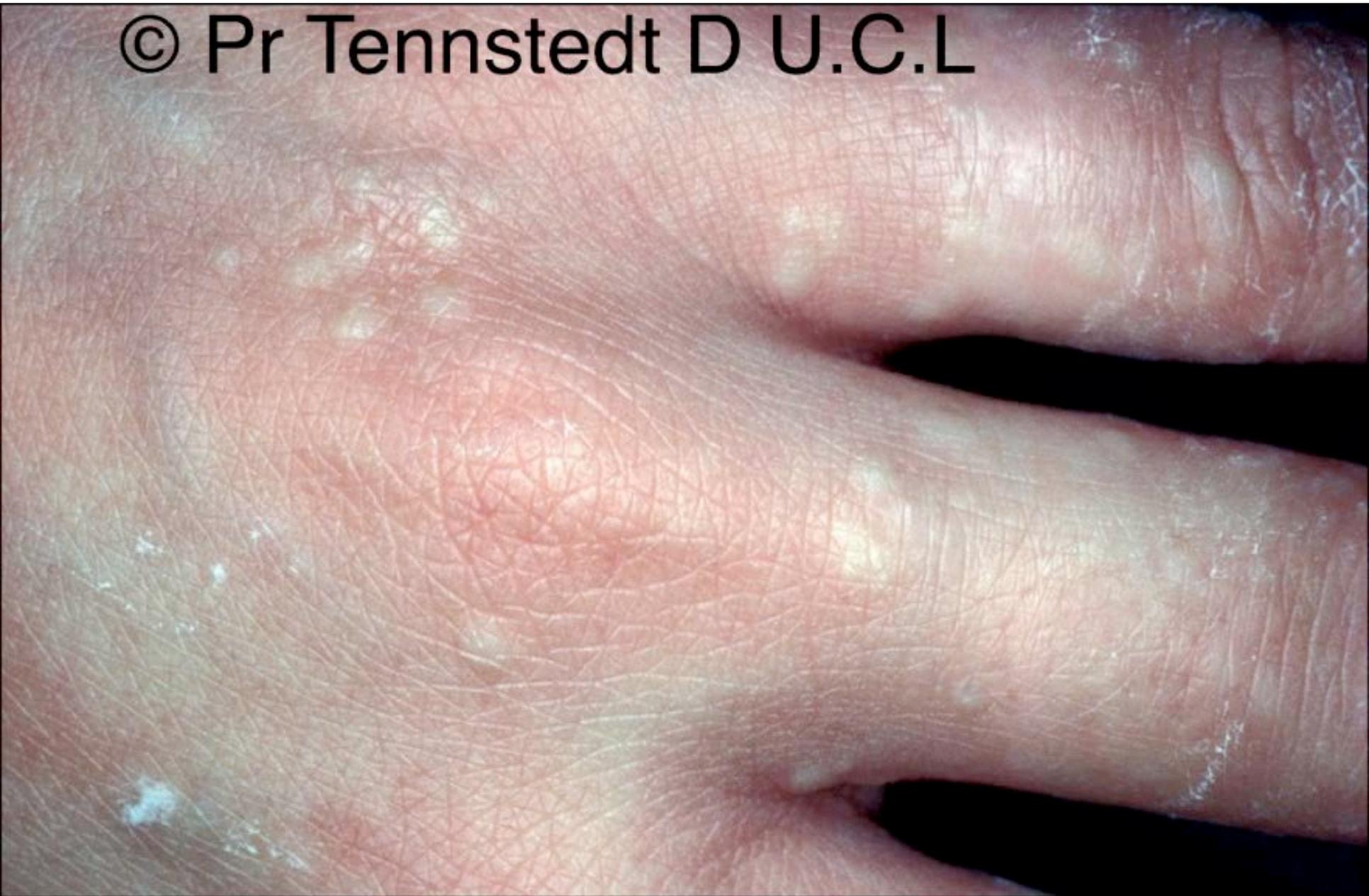
**RUBBER  
ALLERGY**

```
graph TD; A[Immediate : Type I] --> C((RUBBER ALLERGY)); B[Delayed Type IV] --> C;
```

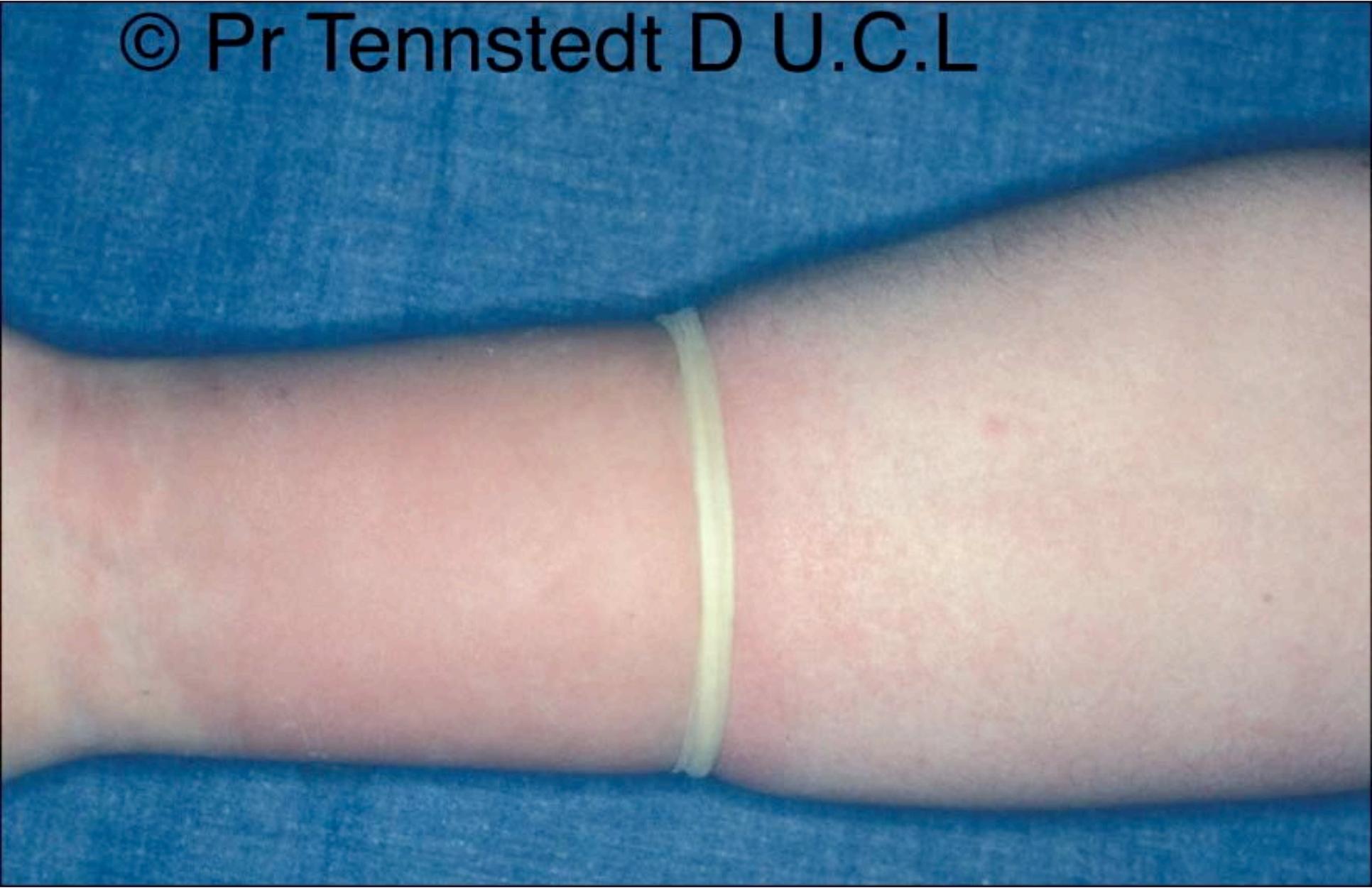
# Natural Rubber latex allergy : immediate reactions



© Pr Tennstedt D U.C.L



© Pr Tennstedt D U.C.L



Diagnosis :  
prick tests  
(and IgE)



**allergie au latex**

# Allergic contact dermatitis (ACD)

- Contact delayed reaction
- eczema



© Dr Crépy MN APHP

© Dr Crépy MN APHP



© Dr Crépy MN APHP



# Patch testing

```
graph TD; A[Patch testing] --- B[European baseline series]; A --- C[Rubber additive series]; A --- D[Gloves];
```

European  
baseline series

Rubber additive  
series

Gloves



## 2) Allergens



# Natural Rubber Latex

Biopolymer : cis-1,4-polyisoprene

Latex : white colloidal suspension (rubber + proteins...)

# Natural rubber latex

- > 200 proteins
- 15 proteins internationally assigned as allergens : Hev b

Allergens

ReTiME

RefArray

Tools

History

Statistics

MyAllergome Shop >

General Information

Sequences

No Natural form

Escherichia coli

ReTiME

## Hev b 15



<b>Entry date</b>	April 27, 2014 12:22 +1GMT
<b>Last update</b>	April 27, 2014 12:24 +1GMT
<b>Allergome Code</b>	11621
<b>Name</b>	Hev b 15
<b>Common Names</b>	<a href="#">Serine protease inhibitor</a>
<b>Biological Function</b>	<a href="#">Serine Protease Inhibitors</a>
<b>Isoforms, Variants, Epitopes</b>	<a href="#">Hev b 15.0101</a>
<b>Links to Molecule Sequences</b>	<a href="#">Hev b 15 - B3FNP9 - UNIPROT</a> , <a href="#">Hev b 15 - Q6XNP7 - UNIPROT</a> , <a href="#">Hev b 15.0101 - W0USW9 - UNIPROT</a>
<b>Sources</b>	<p><a href="#">Euphorbiaceae</a>, <a href="#">Hevea brasiliensis</a>, <a href="#">Latex</a>, <a href="#">Plants</a>, <a href="#">Rubber Tree</a>, <a href="#">Siphonia brasiliensis</a>  <a href="#">... other Source Terms in available Languages -&gt;</a></p> <div style="display: flex; justify-content: space-around;">       </div>
<b>Links to Source Taxonomy</b>	<a href="#">Hev b - 3981 - NCBI</a> , <a href="#">Hev b - 3981 - UniProt</a> , <a href="#">Hev b - 506431 - ITIS</a> , <a href="#">Hev b - Discover Life</a> , <a href="#">Hev b - Wikipedia</a>
<b>Links to Source Images</b>	<a href="#">Hevea brasiliensis on Google images</a>

# Molecular allergens in the diagnosis of latex allergy (*specific IgE, ImmunoCAP*)

Major interest to confirm latex allergy diagnosis :  
Hev b 1, 3, 5, 6.01 et 6.02

Markers of asymptomatic latex sensitivity: Hev b 8,  
(profilin, pan-allergen. pollen-food allergy), carbohydrate  
determinants (CCD)

Hev b 13 : major interest in occupational allergy.  
Currently not available for in vitro diagnosis

# CHEMICAL ADDITIVES

# Rubber vulcanisation additives

Thiurams

dithiocarbamates

benzothiazoles

guanidines

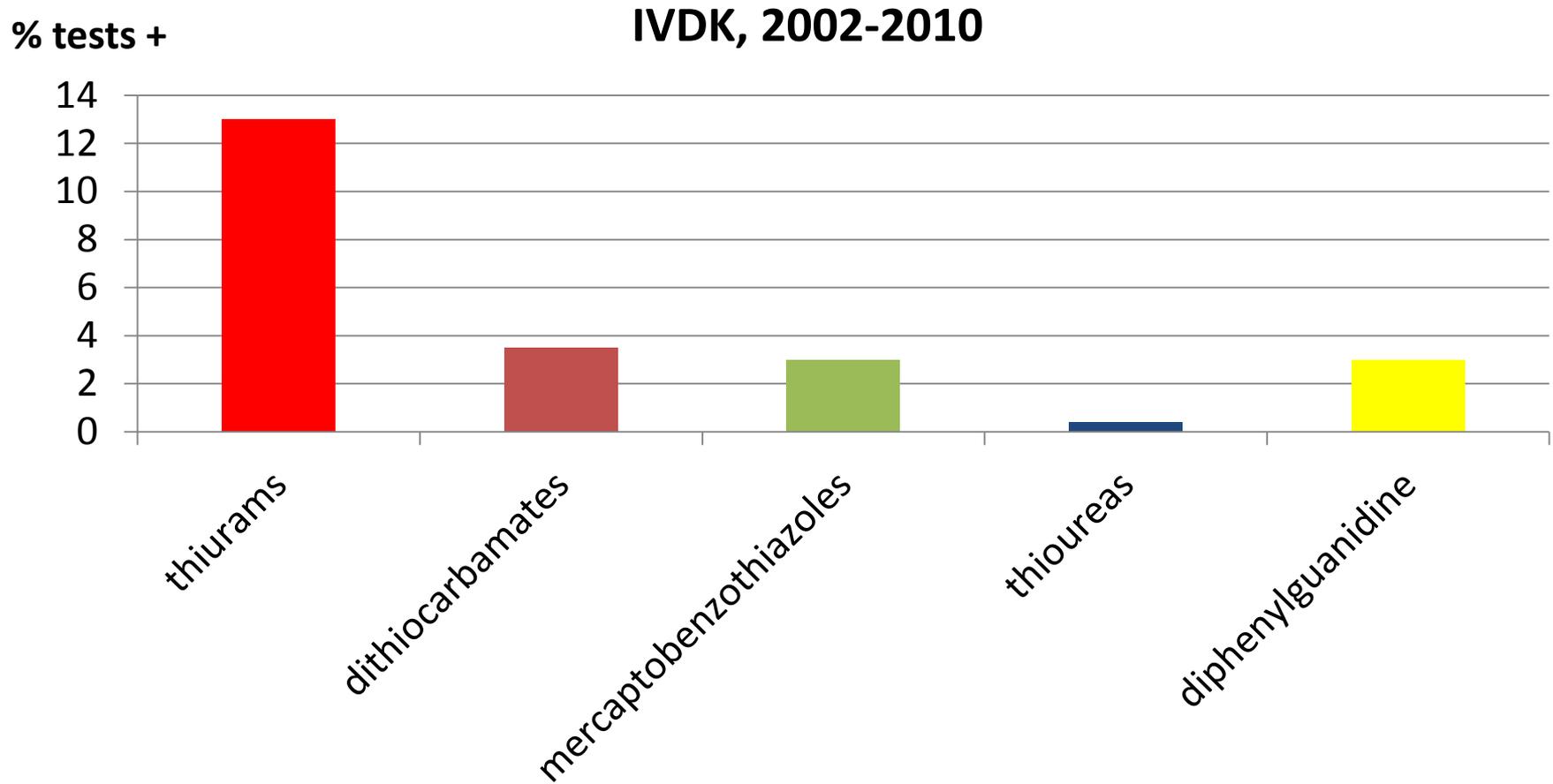
Thioureas

dithiophosphates

Xanthates

(N-(Cyclohexylthio)-*phthalimide*)

# Occupational ACD to rubber gloves



*Geier 2012*

*N= 3448 tested patients for suspected glove allergy*

# thiurames/dithiocarbamates : fast accelerators

## 1. Thiurams :

- Thiurams replaced by dithiocarbamates in gloves
- Thiuram-mix : the best marker of thiuram/dithiocarbamate allergy

## 2. Complex chemical reactions between thiurames and dithiocarbamates :

Thiurams disulfides and dithiocarbamates = redox pair

*(Hansson. CD 2014)*

# Guanidines

- 1,3-diphénylguanidine (gloves...) :  
prevalence ↗



©Crépy MN APHP



© Pr Baeck M. U.C.L

**Thiourea :**  
allergens of  
polychloroprene  
rubber



# Antioxydants

Amines (IPPD..., quinolins)

phenols

dithiocarbamates

Mercaptobenzimidazole  
derivatives

© Dr Crépy MN APHP





© Dr Baeck M. U.C.L



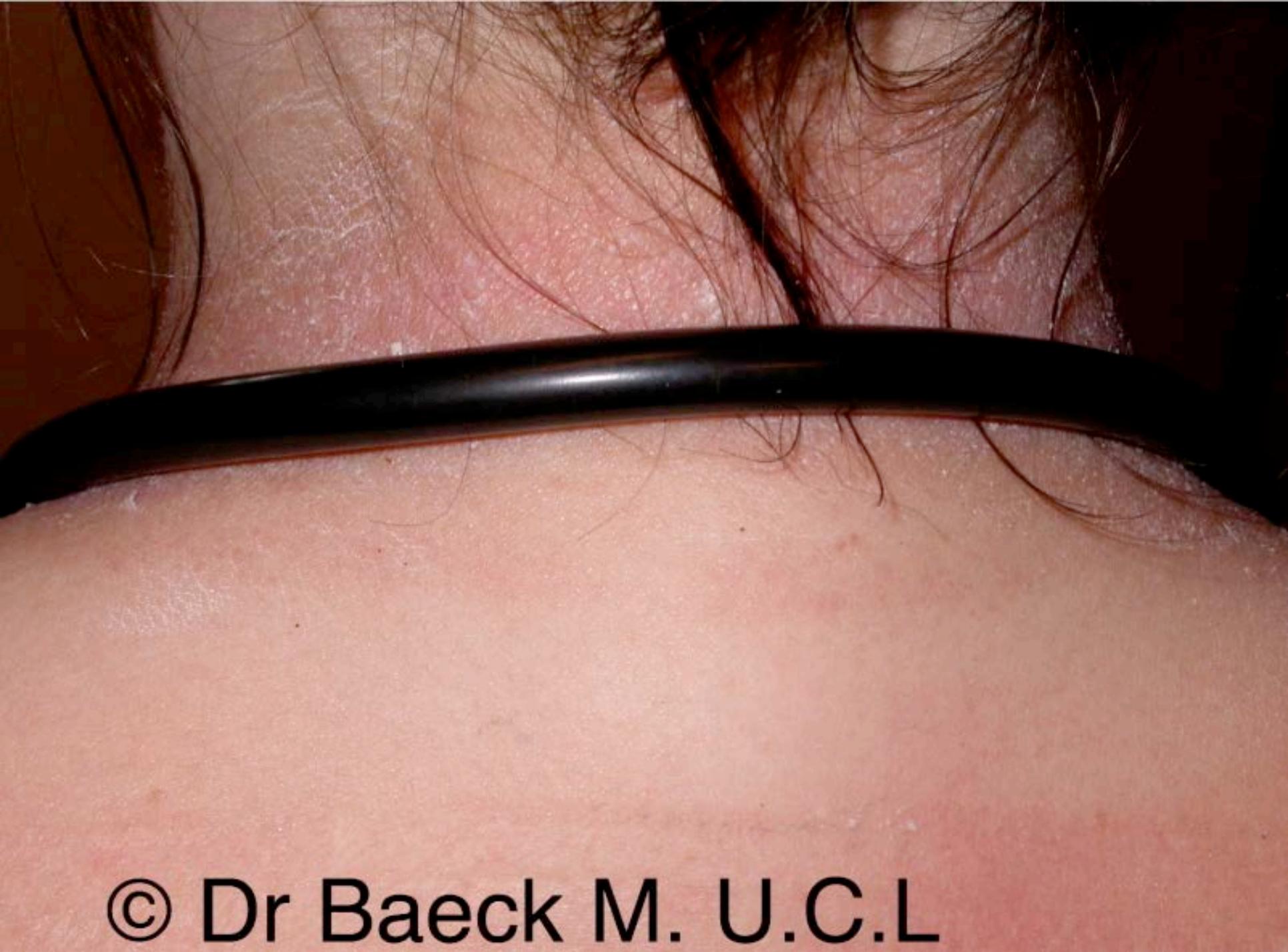
© Dr Baeck M. U.C.L



© Dr Baeck M. U.C.L



© Dr Baeck M. U.C.L



© Dr Baeck M. U.C.L



© Dr Baek M. U.C.L



© Dr Baeck M. U.C.

# Others

plasticiser

surfactants

biocides

Colouring  
agents

others

# Cetylpyridinium chloride : biocide

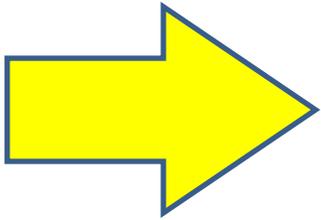
- Eczema (*Ponten, Castelain...*)
- *Donning agent*

### 3) New rubber gloves



# Low protein latex gloves

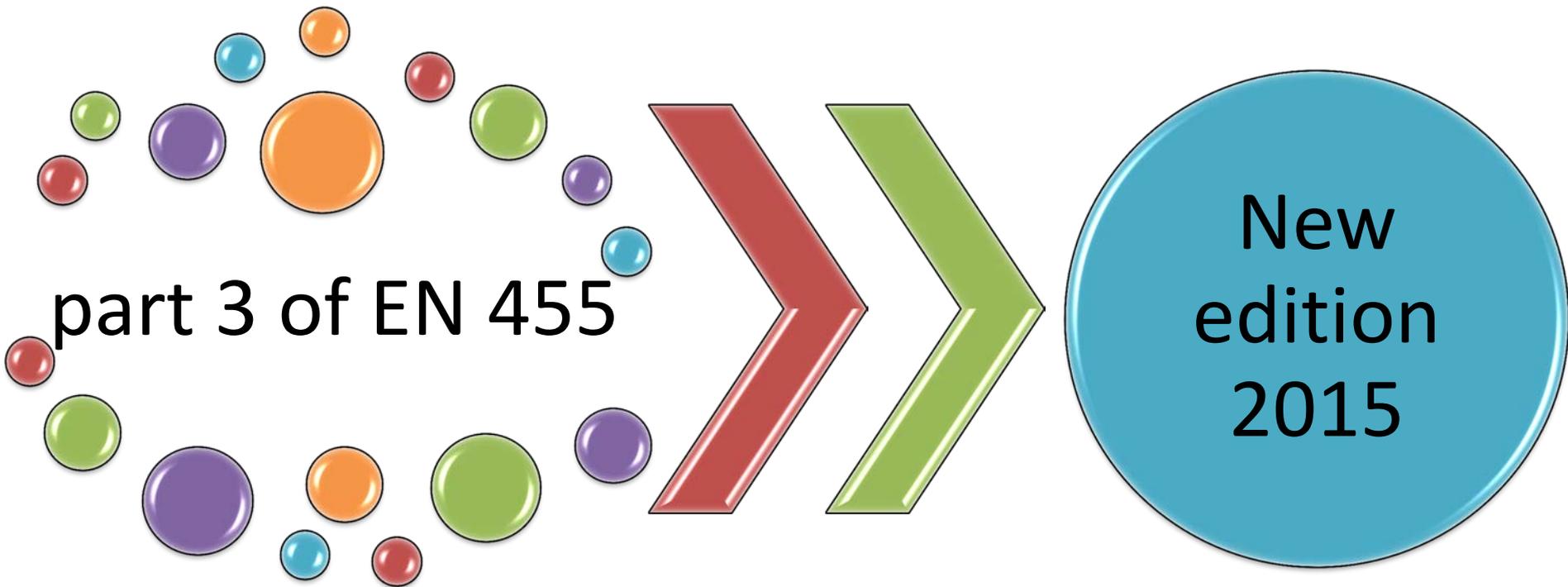
- low-allergens and powder-free medical gloves



successful prevention of NRL-allergy.

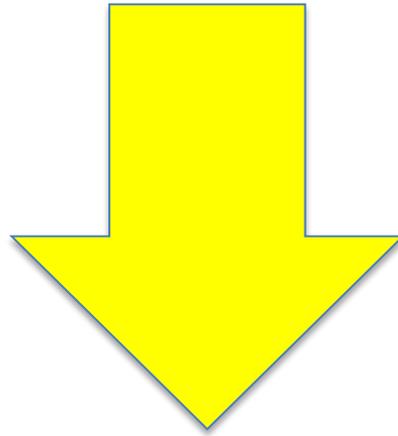
- Significant decline in the occurrence of NRL allergy

# Rubber accelerator-free gloves



# Accelerator-free gloves

may only be labelled as such



**if in no part of the manufacturing process  
accelerators are used**

# Different materials

- Polychloroprene : surgical gloves
- Nitrile : examination gloves
- Polyisoprene with photocrosslinking process : surgical gloves
- Thermoplastic elastomer gloves SEBS : surgical gloves



# Different antimicrobial agents

- didecyldimethylammonium chloride
- Benzalkonium chloride
- chlorhexidine gluconate

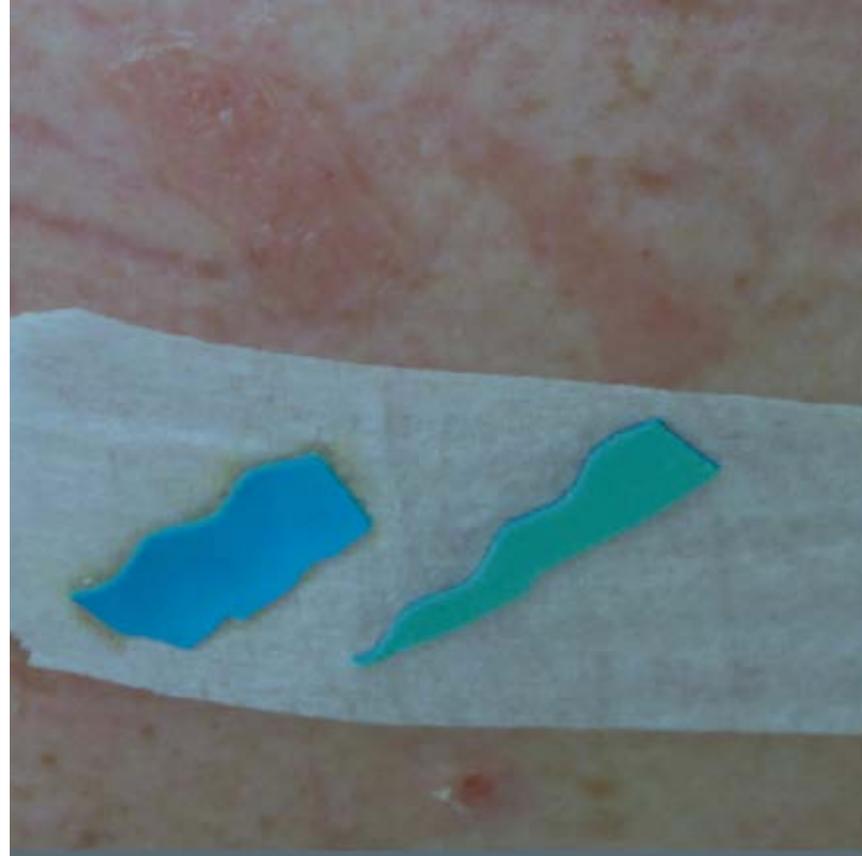
# Moisturizers in gloves !!!

Severe allergic contact dermatitis caused by a rubber glove coated with a moisturizer

**Katrien Vanden Broecke<sup>1</sup>, Erik Zimerson<sup>2</sup>, Magnus Bruze<sup>2</sup> and An Goossens<sup>1</sup>**



- Gloves analysis :
  - fatty alcohols
  - quaternary ammoniums
  
- positive patch test :
  - fatty alcohols
  - quaternary ammoniums



# Guayule (*Parthenium argentatum* Gray)

- alternatives to *hevea brasiliensis* latex



# Conclusion

Latex proteins

Natural rubber latex gloves

Vulcanisation additives

Natural rubber latex gloves

Synthetic gloves

Accelerator-free medical gloves

Biocides in specific surgical glove

New gloves