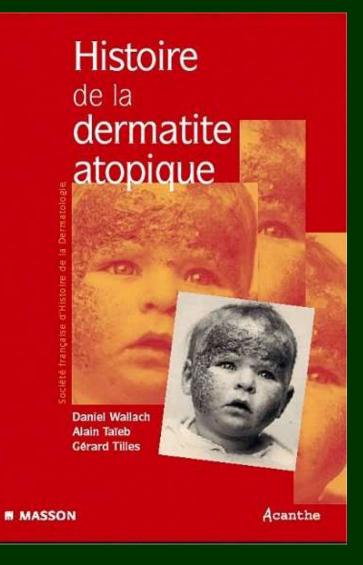


# Atopic dermatitis and dermatological doctrines

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# Can history shed a light on the modern controversies about atopic dermatitis?

- Atopic disorders have been recognized in Ancient times
- Atopic dermatitis was named in 1933
- Before 1933, cases compatible with AD were reported, but with different names
- The descriptions and denominations followed different doctrines
- This is not a surprise, because even now, there is no consensus on the nosology of atopic dermatitis
- The current lack of consensus can be interpreted as the consequence of divergent medical doctrines, i.e allergological vs. dermatological

# Children like this one have probably been observed since Antiquity.



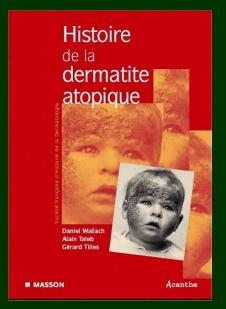
# **But what did the physicians see?**

- An oozing disease
- A disease starting on the head and scalp
- A disease of suckling infants
- A vesicular dermatitis
- A papular, pruritic dermatitis
- Pruritus
- Skin lesions secondary to pruritus
- An allergic disease
- A hereditary, hyper-IgE disease
- A digestive disease
- The result of a conflict between the child and his/her mother



# What do we learn from history?

• Key message :



when physicians see patients,

the disease they describe,

the way they name it,

and the way they treat it,

mainly depend upon their medical doctrine

# **Atopic dermatitis : three key dates**

- 1796 : prurigo/eczema/...
- 1892 : diathetic prurigo
- 1933 : atopic dermatitis



# **Atopic dermatitis : four periods**

**Before 1800 : ancient (hippocratic) medicine** 

1796 : eczema / prurigo/...

Nineteenth century : willanist clinical approach

• 1892 : diathetic prurigo

**Diathesis and immunology** 

• 1933 : atopic dermatitis

What is atopic dermatitis?



## **Humoralist doctrine**

• 1796 : eczema / prurigo/...

Willanist doctrine

• 1892 : diathetic prurigo

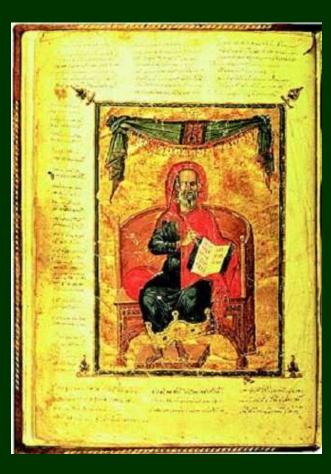
**Diathetic doctrine** 

• 1933 : atopic dermatitis

Allergo / Immunological doctrine

# The four periods of atopic dermatitis history

# First period : before 1800, skin diseases according to ancient medicine



### Atopic dermatitis in ancient medicine : Mercurialis

 Girolamo Mercurialis (1530-1606), first author of a book
 specifically « dermatological » :

De morbis cutaneis et omnibus corporis humani excrementis (1572)

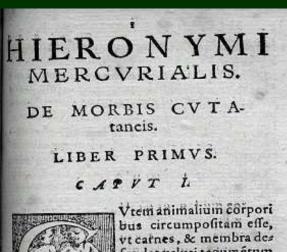


#### De morbis cutaneis et omnibus corporis humani excrementis (1572)

#### Livre I : Dermatoses de la tête

Ch 1 : introduction, généralités Ch 2 : classification des maladies de la peau de la tête Ch 3 : chutes de cheveux Ch 4 : alopécies Ch 5 : calvitie Ch 6 : canitie Ch 7 : pédiculose Ch 8 : porrigo (pityriasis) Ch 9 : achore et favus Ch 10 : teigne Ch 11 : vésiculo-pustules, ulcères superficiels, sycosis, exanthèmes Livre II : Dermatoses du corps Ch 1 : généralités sur la classification

Ch 1 : généralités sur la classification Ch 2 : dépigmentations, troubles pigmentaires Ch 3 : prurit Ch 4 : gale Ch 5 : lèpre Ch 6 : lichens





Vtem animalium corport bus circumpolitam effe, vt carnes, & membra des fendat veluti tegumétum quoddám, clarifsimè conftat apud omnes medicos. Quanquam Hippocrates in libro de Ofsibus, alium

finem habuiffe Naturam in fabricanda cute di xiffe videatur: vbi fcribit, Eam effe veluti quod dam membrorum ligamentum, quandoquidé cum membra fint omnia veluti inter fe fepara n,opus fuit communi quodam ligamine conuinciri, quod quidem fuit cutis. Vnde fit, vt iuremerito Diuinus Plato in Timæo cutim naffulz pifcatoriæ adfimilauerit, quod etiam adnotauit Galenus in Commentarijs in Timæum. Cum itaque fit hic folus cutis vfus, cum fi nullam aliam actionem vtilem habeat, fed folu txcrementorum fit receptaculú, vt docuit Ga

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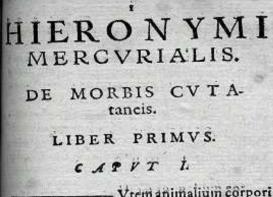
#### Ch 9 : achore et favus

Ch 10 : teigne

Ch 11 : vésiculo-pustules, ulcères superficiels, sycosis, exanthèmes

#### Livre II : Dermatoses du corps

Ch 1 : généralités sur la classification Ch 2 : dépigmentations, troubles pigmentaires Ch 3 : prurit Ch 4 : gale Ch 5 : lèpre Ch 6 : lichens



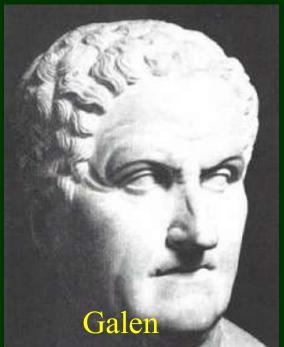


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# Mercurialis' humoralist doctrine

- Skin disorders were viewed as a route of elimination of humors
- Phlegmatic diseases were more prevalent in winter
- Classification : head body
- Among scalp dermatoses :
  - Achores : scalp oozing conditions
    - Head ulcer
    - Mucous tinea



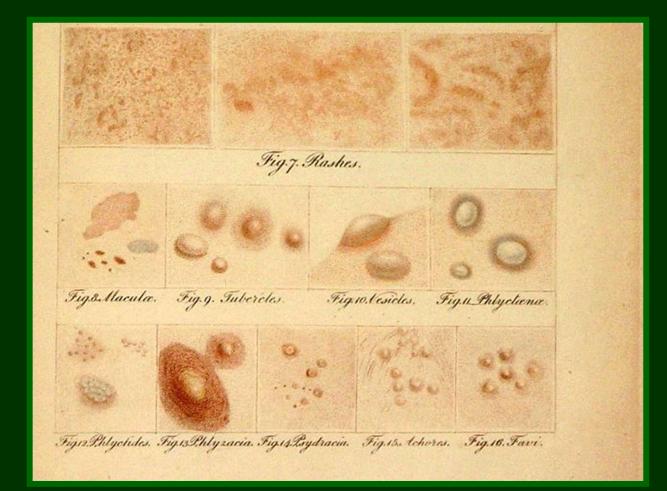
# **Achores according to Mercurialis**

- Lesions of the scalp, from which humors eliminate
- Background :
  - Infants (excretions accumulate in utero)
- Etiology : the « infectious » milk from the mother / wet nurse
- Diagnosis :

   Pruritus, then lesions and oozing
   Color and taste characteristic
- Treatment
  - The wet nurse's diet
  - Very few drugs

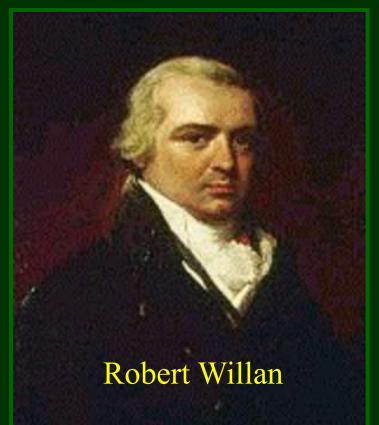


# Second period : willanism



# Willan-Bateman : « Birth of the clinics » in dermatology

- Robert Willan (1757-1812) :
  - Physician of a public dispensary
  - On cutaneous diseases (1798-1808)
- Thomas Bateman (1778-1821)
  - Faithful and talented friend, pupil, follower
  - A practical synopsis ....
  - Delineations ...



#### Willan-Bateman :

# after Plenck, a classification on a semiological basis

Order I : Papules	<b>Order IV : Bullae</b>		Order VII : Tubercles	
Strophulus		Pemphigus		Phyma
Lichen		Pompholyx		Warts
Prurigo	<b>Order V : Pustules</b>			Molluscum
		Impetigo		Vitiligo
Order II : Squamae		Porrigo		Acne
Lepra		Ecthyma		Sycosis
Psoriasis		Smallpox		Lupus
Pityriasis		Scabies		Elephantiasis
Ichtyosis	<b>Order VI : Vesicles</b>			Framboesia
		Varicella		Tamotesia
<b>Order III : Exanthemas</b>		Vaccinia	<b>Order VIII : Macules</b>	
Rubeola		Herpes	Order vin Macules	Ephelis
Scarlet fever		Rupia		Naevus
Urticaria		Miliaria		INACVUS
Roseola		Eczema		
Purpura		Aphtae		
Erythème		1		
Erysipela				

# Atopic dermatitis in Willan/Bateman's works

• Where is it?

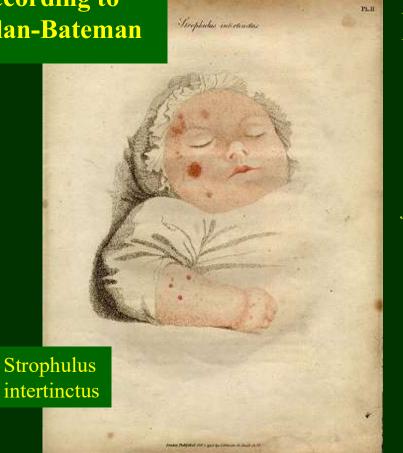
- Was « atopic dermatitis » described as an autonomous entity?
- If not, why did not Willan/Bateman identify atopic dermatitis?

### Willan-Bateman :

# descriptions compatible with atopic dermatitis

Order I : Papules	Order IV : Bullae		Order VII : Tubercles	
Strophulus		Pemphigus		Phyma
Lichen		Pompholyx		Warts
Prurigo	<b>Order V : Pustules</b>			Molluscum
		Impetigo		Vitiligo
<b>Order II : Squamae</b>		Porrigo		Acne
Lepra		Ecthyma		Sycosis
Psoriasis		Smallpox		Lupus
Pityriasis		Scabies		Elephantiasis
Ichtyosis	<b>Order VI : Vesicles</b>			Framboesia
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Rubeola		Herpes		Ephelis
Scarlet fever		Rupia		Naevus
Urticaria		Miliaria		11401405
Roseola		Eczema		
Purpura		Aphtae		
Erythème				
Erysipela				

#### Strophulus according to Willan-Bateman



*"the Strophulus is a papulous eruption, peculiar to infants, and exhibiting a variety of forms, ...* 

« Strophulus confertus : An eruption of numerous papules, varying in their size, appears on different parts of the body in infants, during dentition ....and has then been denominated the Tooth Rash"

"About the fourth or fifth month after birth, an eruption of this kind takes place, in most infants, on the cheeks, and sides of the nose, extending sometimes to the forehead and arms, sometimes, though less frequently; to the trunk of the body"



#### Lichens according to Willan-Bateman

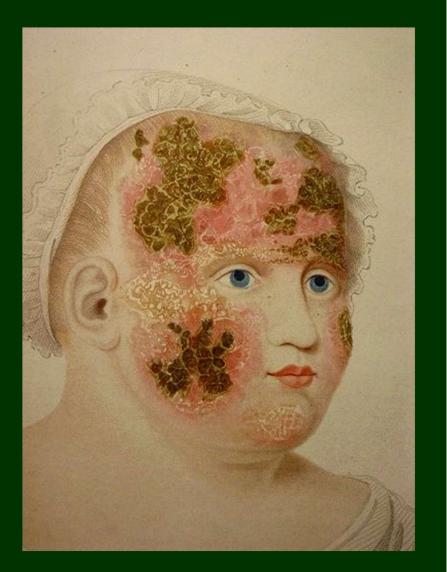
• "an extensive eruption of Papulae, affecting adults, connected with internal disorder, usually terminating in Scurf ; recurrent ; non contagious".

• Seven forms



#### **Porrigo larvalis willani**

- In the order of the pustules
- "..., or Crusta lactea of authors, is almost exclusively a disease of infancy. It commonly appears first on the forehead and cheeks, in an eruption of minute and whitish achores, which are crowded together, upon a red surface. These pustules soon break, and discharge a viscid fluid, which concretes into thin yellowish or greenish scabs ".
- Porrigo # tinea # favus # achore = an oozing / crusty scalp dermatosis
- Pruritus is important



#### **Eczema according to Willan-Bateman**

• Sixth gender of the Order of vesicular dermatoses.

«The eczema is characterized by an eruption of small vesicles, on various parts of the skin, usually set close or crowded together, with little or no inflammation round their bases, and unattended by fever. It is not contagious. ... ».

 « usually secondary to an irritation, of external or internal origin, and may be produced by a great variety of irritants, in persons whose skin is constitutionally very irriatble».





# Atopic dermatitis in Willan/Bateman's works

• Where is it?

Not among eczemas, but among varieties of papular/ pustular diseases

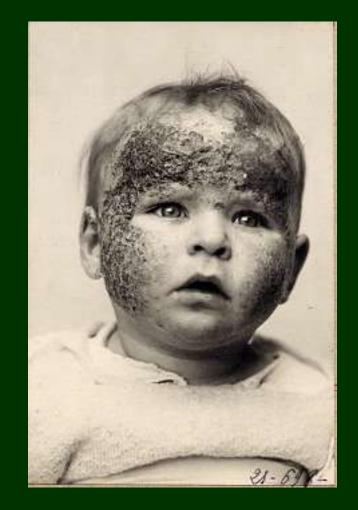
• Was « atopic dermatitis » described as an autonomous entity?

No

• If not, why did not Willan/Bateman identify atopic dermatitis?

#### Why did willanists fail to identify atopic dermatitis?

- Because AD does not have only one primary (elementary) lesion
  - At a given time
  - During the course of the disease



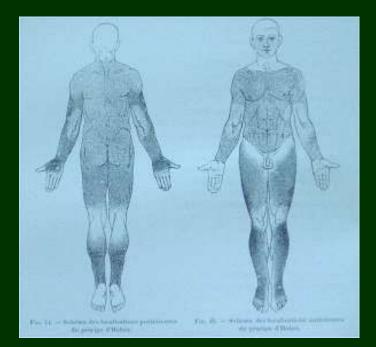
Third period : from papular prurigo (Hebra) to diathetic prurigo (Besnier)





# Hebra's prurigo

« In all cases, the first phenomenon consists in the presence of subepidermal papules ... »

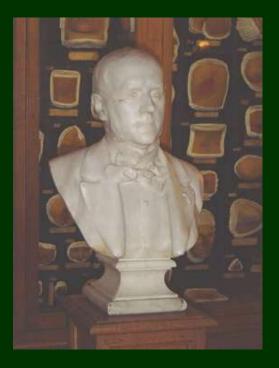




Rurigo agria sen forex mihi.

## **1892 : Ernest Besnier** « gets free from the willanist tyranny »

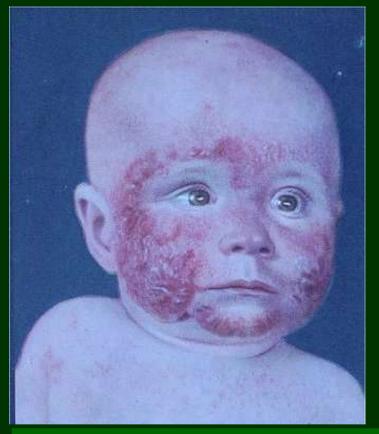
- Besnier Ernest. Première note et observations préliminaires pour servir d'introduction à l'étude des prurigos diathésiques Ann Dermatol Syphil. Troisième série, tome III, 1892 : 634-648.
- « Their first symptom and main symptom is pruritus »
- « In many cases, the pathological process leaves the skin, to have visceral localisations including emphysema, bronchial asthma, hay fever, end more rarely digestive troubles »



# Fourth period : immunology meets eczema



#### A conceptual challenge : to link infantile eczema and chronic prurigo



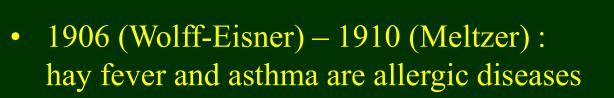
Infantile, constitutional eczema

#### Hebra's prurigo agria

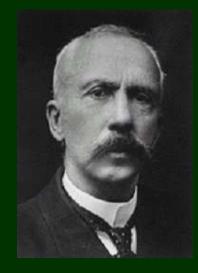


#### Advances in immunological hypersensitivities

- 1902 : Richet and Portier : anaphylaxis, an experimental concept.
- 1906 : von Pirquet : allergy : a change in clinical reactivity



• 1916 (Blackfan) : infantile eczema and allergy to food proteins





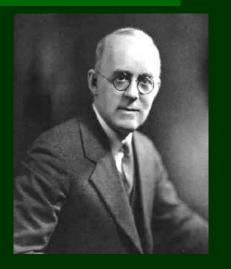
# Classification of hypersensitivities, Coca and Cooke, 1923

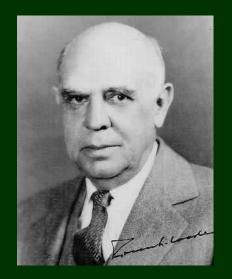
# Hypersensitivity

- Normal (usual)
  - Dermatitis venenata (contact eczema)
  - Serum sickness

## – Abnormal

- Anaphylaxis
- Hypersensitivity of infections (tuberculosis, others)
- Atopy : hereditary hypersensitivity to environmental antigens

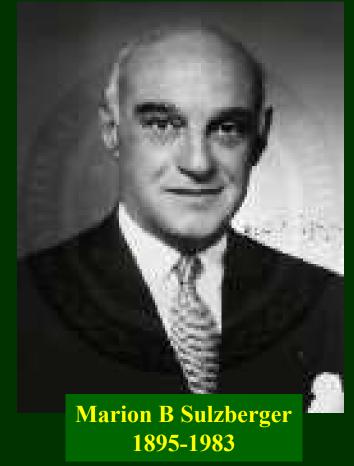




# Wise and Sulzberger, 1933 : « This is probably best called atopic dermatitis »

### Among previous denominations

- Eczematoïd exsudative, infantile eczema, asthma-eczema, asthma-prurigo, flexural eczema, Besnier's prurigo, neurodermatitis, lichenifications, generalized pruritus with lichenification, ...
- Wise and Sulzberger pick up a group of chronic eczemas with positive skin tests,
- Which « is probably best called atopic dermatitis »,
- And for which some criteria may be defined



# The 1933 definition of atopic dermatitis

(Wise F, Sulzberger MB. 1933 Year Book of Dermatology and Syphilology ; 38-39.)

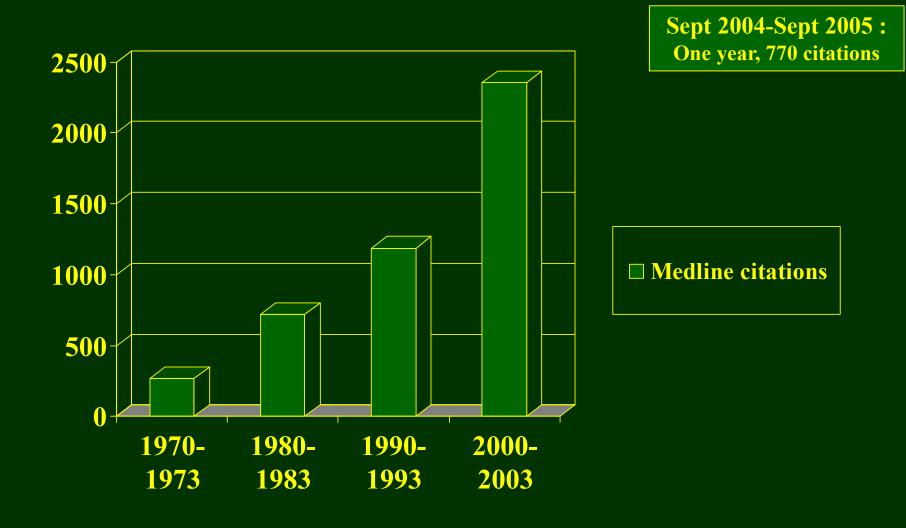
- 1 Atopic family history ;
- 2 Antecedent infantile eczema ;
- 3 Localization in antecubital and popliteal spaces, the anterior portions of the neck and chest and the face, particularly the eyelids ;
- 4 A greyish or brownish coloration of the skin
- 5 The absence of true vesicles, clinically and histologically
- 6 Vasomotor instability or irritability
- 7 Negativity of patch tests to contact irritants (particularly adhesive)
- 8 Many positive reactions of immediate wheal type to scratch or intradermal testing
- 9 The presence of many reagins in the blood serum.

# Since 1933, an unsolved question : what is really atopic dermatitis?





## **1933-2001 : a growing interest for atopic dermatitis**



# Atopic dermatitis, 1933-2001 : progress

- Unified terminology
- Validated diagnostic criteria
  - Hanifin / Rajka
  - UK Working Party



- A clinical scoring for severity, widely accepted
   SCORAD
- An efficient therapy
  - Topical steroids

#### Atopic dermatitis, 1933-2001 : unsolved questions

- Terminology
  - Dermatitis or eczema?
  - intrinsic, atopiform, non atopic, entopic, ...
- Role of IgE-mediated allergy
  - For the diagnosis
  - For understanding the pathophysiology
  - For the management
    - Diet, food eliminations, probiotics, hyposensitizations, ...
- Confidence in topical steroid therapy
- And in rational medicine



# The 2005 debate

• Is atopic dermatitis defined by clinical criteria? (Hanifin-Rajka's, UKWP's)?

Dermatological doctrine : AD is a skin disorder IgE are not part of the definition and are not routinely assayed

• Or is atopy confined to IgE sensitization?

Immuno-allergological doctrine : IgE define atopy AD is the cutaneous component of the atopic disease

# **Conclusion : historical conceptions of atopic dermatitis**

- A reflection of internal disorders
- A skin disease
- A diathesis
- An allergological disorder



Will the 4<sup>th</sup> Georg Rajka ISAD resolve all the questions?

> 4th Georg Rajka International Symposium on Atopic Dermatitis (SAC) ARCACHON, FRANCE 15-17 SEPT 2005

# If not, a suggested reading (in French)

# Histoire de la dermatite atopique

Société française d'Histoire de la Dermatol-

Thank you for your attention

Daniel Wallach Alain Taïeb Gérard Tilles

**MASSON** 

Acanthe