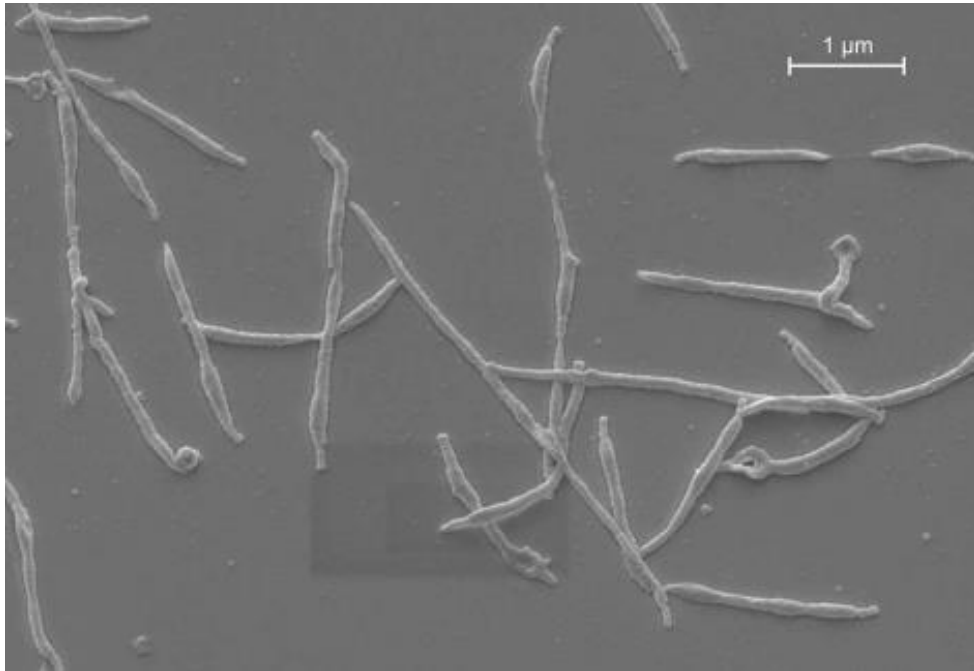


# L'infection à Mycoplasme en pédiatrie

Philippe Minodier, *Hôpital Nord, APHM, Marseille*

Ania Carsin, *Hôpital St Joseph, Marseille*

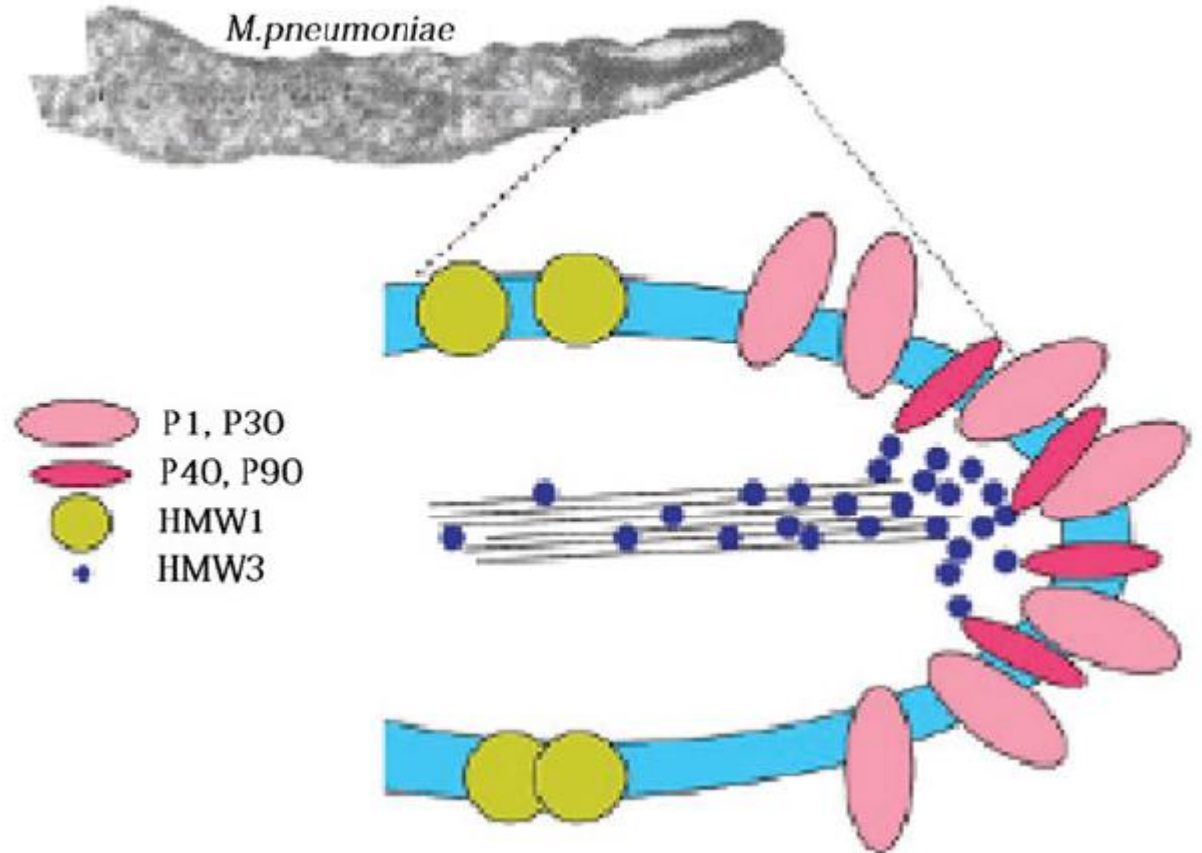
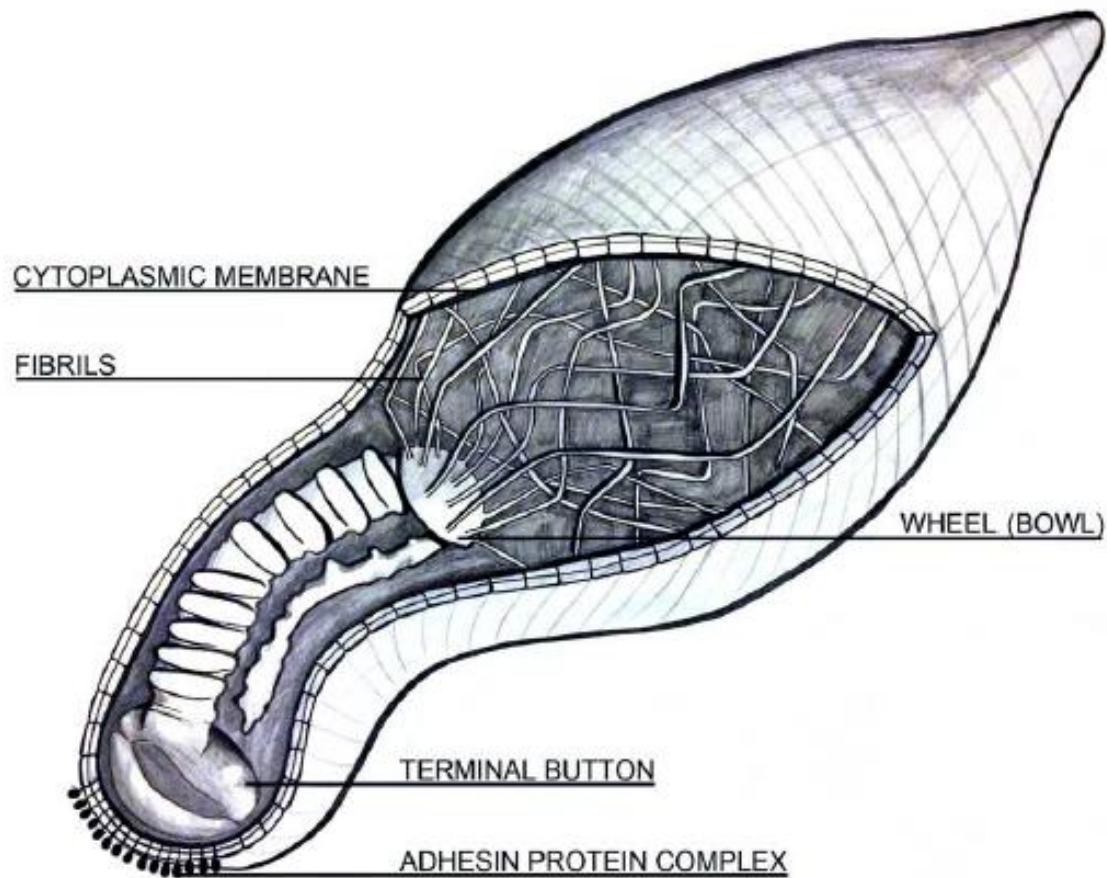
# Mycoplasma pneumoniae



Pas de peptidoglycane  
Pas d'efficacité des  $\beta$ -lactamines

- Découvert en 1944 par **Eaton**
- Initialement considéré comme un virus
- 1950-1960 : déclenche des manifestations respiratoires hautes et basses chez des volontaires
- 1<sup>ère</sup> culture sur milieu acellulaire en 1962
- Taille : **1-2  $\mu\text{m}$  x 0,1-0,2  $\mu\text{m}$**
- Génome (1996) : 816 394 bp, **687 gènes**

# L'organelle d'attachement



# Pathogénicité

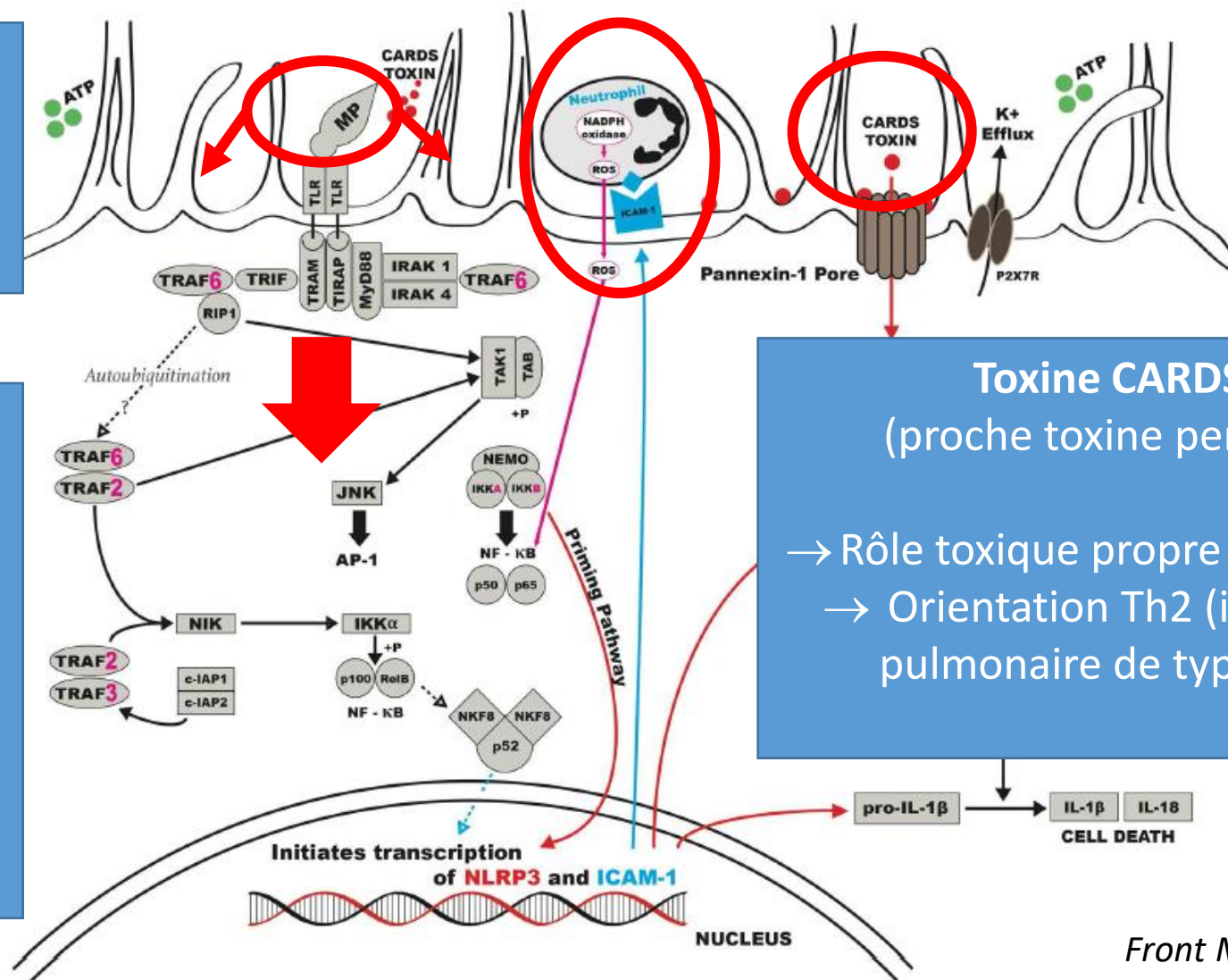
Production ions peroxydes  
et superoxydes

→ Effet toxique direct

Activation de la cascade  
inflammatoire

Production ICAM-1  
(adhésion Mφ)

→ Effet toxique indirect



Toxine CARD6 +++

(proche toxine pertussique)

→ Rôle toxique propre (vacuolisation)

→ Orientation Th2 (inflammation  
pulmonaire de type allergique)

pro-IL-1β → IL-1β IL-18  
CELL DEATH

# Données épidémiologiques

- Transmission par **gouttelettes respiratoires**
- **2 sous-types** avec variants (recombinaisons entre séquences répétées RepMP2/3 et RepMP4 des gènes codant pour protéine P1)
- **Immunité peu durable** : réinfections, récurrences
- Évolution cyclique tous les 3-7 ans
- **Prévalence difficile à établir** :
  - Selon la technique de dépistage : PCR, sérologie
  - Selon la gravité : beaucoup de patients traités en ambulatoire
  - Selon l'âge : rare avant 4 ans, pathologie du grand enfant et de l'adolescent
  - Portage asymptomatique

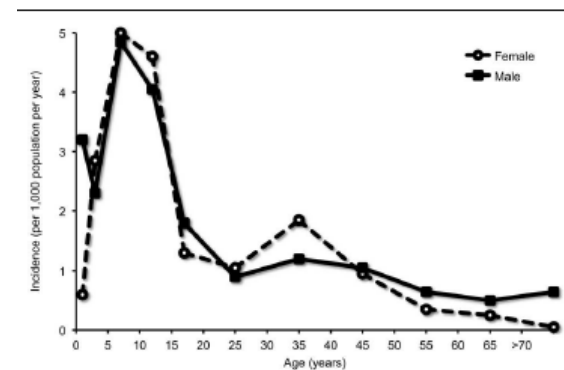


FIGURE 2 | Detection of *M. pneumoniae* in community-acquired pneumonia (CAP) according to age group. Infection was diagnosed by culture of respiratory specimens and/or a fourfold titer rise in complement fixation test (CFT). Adapted with permission from Foy et al. (1979).

# Modalités du diagnostic

# Diagnostic : sérologie

- **IgM (IgA)** : 1<sup>ère</sup> semaine de la maladie, pic à 3 semaines
- **IgG** : séroconversion à 3-8 semaines
  
- Techniques : **enzyme immunoassays** (EIA) les plus évalués
  
- **Inconvénients** :
  - Manque de sensibilité
  - Faux positifs entre différentes espèces de *Mycoplasma*
  - Délai de 2 semaines ou plus avant séroconversion
  - Faux négatifs chez immunodéprimés

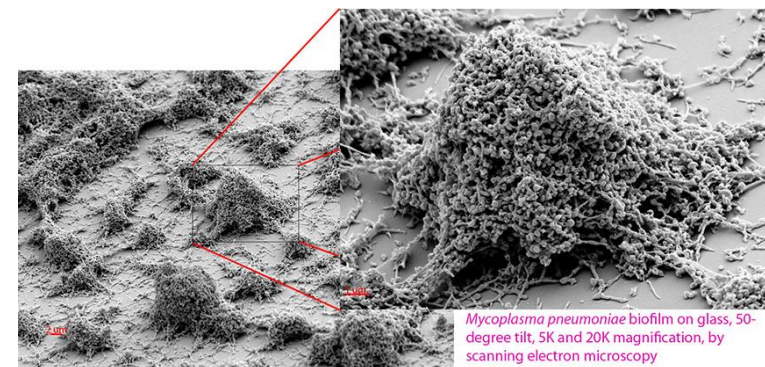
# Diagnostic : biologie moléculaire (PCR)

- **Différentes techniques**

- PCR conventionnelle
- Nested PCR (réamplification par un 2<sup>ème</sup> jeu de primers) : ↑ Sensibilité
- PCR quantitative en temps réel : distingue les sous-types, différencie (?) infection aiguë et le portage
- PCR multiplex : permet diagnostic de plusieurs germes (pathologie respiratoire)
- PCR LAMP : technique plus simple... mais moins sensible

- **Différentes cibles**

- ARN 16S
- Protéine d'adhésion P1
- Gènes de l'opéron ATPase
- Toxine CARDS



| Method  | Test                                  | Target/antigen  | Antibodies                               | Specimen(s)   | Performance  | Value          | Comments  |
|---|---------------------------------------|---|--|---|--|----------------|---|
| Direct identification of <i>M. pneumoniae</i>           | Polymerase chain reaction (PCR)       | Different target genes (e.g., P1 gene, 16S rDNA, 16S rRNA, RepMP elements etc.) | –  | Respiratory specimen<br>Cerebrospinal fluid (CSF)<br>Other bodily fluids or tissues | High sensitivity, high specificity                   | RD             | <ul style="list-style-type: none"> <li>- Validation and standardization required for routine diagnostic (Loens et al., 2010);</li> <li>- Epidemiological differentiation of clinical strains on the basis of differences in the P1 gene by PCR (Spuesens et al., 2009) or in the number of repetitive sequences at a given genomic locus by multiple-locus variable-number tandem repeat analysis (MLVA) (Chalker et al., 2015).</li> </ul> |
| Non-specific serological tests for <i>M. pneumoniae</i> | Culture                               | –   | –  | Respiratory specimen  | Low sensitivity, high specificity                    | AD             | <ul style="list-style-type: none"> <li>- Special enriched broth or agar media;</li> <li>- Isolation takes up to 21 days.</li> </ul>   |
|   | Cold agglutinin test (“bedside test”) | Erythrocytes (I antigen)  | Cold agglutinins (IgM)                   | Serum   | Low sensitivity, low specificity                     | – <sup>1</sup> | <ul style="list-style-type: none"> <li>- Cold agglutinins target the I antigen of erythrocytes;</li> <li>- Positive in only about 50% and in the first week of symptoms;</li> <li>- Less well studied in children;</li> <li>- Cross-reactivity with other pathogens and non-infectious diseases.</li> </ul>   |
| Specific serological tests for <i>M. pneumoniae</i>     | Complement fixation test (CFT)        | Crude antigen extract with glycolipids and/or proteins                          | Igs (no discrimination between isotypes) | Serum   | Sensitivity and specificity comparable to EIA        | – <sup>1</sup> | <ul style="list-style-type: none"> <li>- Positive criteria: fourfold titer increase between acute and convalescent sera or single titer <math>\geq 1:32</math>;</li> <li>- Cross-reactivity with other pathogens and non-infectious diseases.</li> </ul>  |
|   | Particle agglutination assay (PA)     |   | IgM and IgG simultaneously               |   |  | – <sup>1</sup> |   |
|   | Enzyme immunoassay (EIA)              | Proteins (e.g., adhesion protein P1) and/or glycolipids                         | IgM, IgG, IgA                            | Serum<br>CSF <sup>2</sup>   | Moderate-high sensitivity, Moderate-high specificity | RD             | <ul style="list-style-type: none"> <li>- The sensitivity depends on the time point of the first serum and on the availability of paired sera (for seroconversion and/or rise in titer);</li> <li>- “Gold standard”: fourfold titer increase as measured in paired sera.</li> </ul>  |
|   | Immunoblotting                        |   |  |   | High sensitivity, high specificity                   | AD             | <ul style="list-style-type: none"> <li>- Confirmatory assay (Dumke et al., 2012).</li> </ul>  |
|   | Immunofluorescent assay (IFA)         |   |  |   | Less sensitive and less specific than EIA            | AD             | <ul style="list-style-type: none"> <li>- Subjective interpretation.</li> </ul>  |

# Le portage sain

# La problématique du portage (sain)

## Carriage of *Mycoplasma pneumoniae* in the Upper Respiratory Tract of Symptomatic and Asymptomatic Children: An Observational Study

- Rotterdam (Pays Bas)
- **Enfant 3 mois – 16 ans**, sans pathologie chronique
- 2008-2011
- **Asymptomatiques (n = 405)**  
**ou infection des VA (n = 321)**
- rt-PCR pharynx, naso-pharynx + sérologie
- **Rt-PCR positive**
  - Asymptomatiques : 21,2%
  - Symptomatiques : 16,2%

}  $p = 0,11$
- **Chez symptomatiques :**
  - VAS : 15,9%
  - VAI : 15,6%

}  $p = 0,85$

# La problématique du portage (sain)

*Plos Med 2013;10:e1001444*

**Table 2.** Results from the multiple logistic regression analysis for a positive *M. pneumoniae* PCR result in the asymptomatic group.

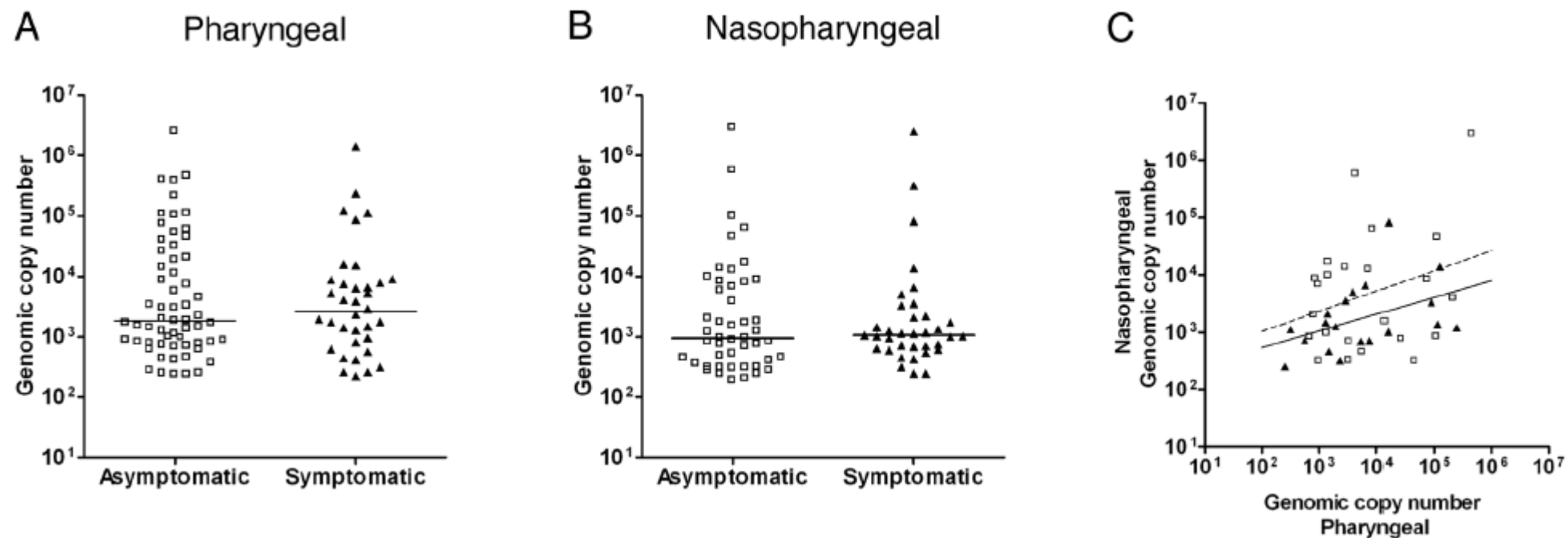
| Variable                               | Odds Ratio (95% CI) | p-Value          |
|--|---------------------|------------------|
| Age ( $\geq 5$ y)                      | 0.96 (0.47–1.96)    | 0.91             |
| Gender (female)                        | 1.44 (0.81–2.56)    | 0.22             |
| Season                                 |                     | <0.001 (overall) |
| Spring versus winter                   | 0.81 (0.33–1.97)    | 0.64             |
| Summer versus winter                   | 7.43 (3.09–17.85)   | <0.001           |
| Autumn versus winter                   | 2.90 (1.26–6.70)    | 0.01             |
| Year (2009 versus 2010 and 2011)       | 3.31 (1.75–6.27)    | <0.001           |
| Family size ( $\geq 5$ family members) | 1.55 (0.77–3.13)    | 0.22             |
| Smoking (active or passive)            | 0.70 (0.38–1.26)    | 0.23             |
| Presence or history of wheezing        | 2.30 (0.69–7.61)    | 0.17             |
| Day-care attendance                    | 0.82 (0.38–1.77)    | 0.62             |
| RTI prior to enrollment                | 0.95 (0.50–1.79)    | 0.86             |
| RTI post-enrollment                    | 0.59 (0.29–1.20)    | 0.15             |

**Table 3.** Results from the multiple logistic regression analysis for a positive *M. pneumoniae* PCR result in the symptomatic group.

| Variable                               | Odds Ratio (95% CI) | p-Value        |
|--|---------------------|----------------|
| Age ( $\geq 5$ y)                      | 1.56 (0.60–4.02)    | 0.36           |
| Gender (female)                        | 0.93 (0.46–1.87)    | 0.84           |
| Season                                 |                     | 0.87 (overall) |
| Spring versus winter                   | 0.80 (0.35–1.81)    | 0.59           |
| Summer versus winter                   | 0.54 (0.11–2.67)    | 0.45           |
| Autumn versus winter                   | 0.85 (0.39–2.16)    | 0.74           |
| Year (2009 versus 2010 and 2011)       | 5.80 (1.94–17.35)   | 0.002          |
| Family size ( $\geq 5$ family members) | 1.63 (0.74–3.61)    | 0.23           |
| Smoking (active or passive)            | 0.61 (0.30–1.23)    | 0.17           |
| Presence or history of wheezing        | 1.96 (0.93–4.13)    | 0.08           |
| Day-care attendance                    | 0.84 (0.37–1.89)    | 0.67           |
| Diagnosis (lower RTI)                  | 1.05 (0.46–2.42)    | 0.91           |
| Hospitalization                        | 1.54 (0.52–4.60)    | 0.44           |

# Portage vs infection : type de prélèvement

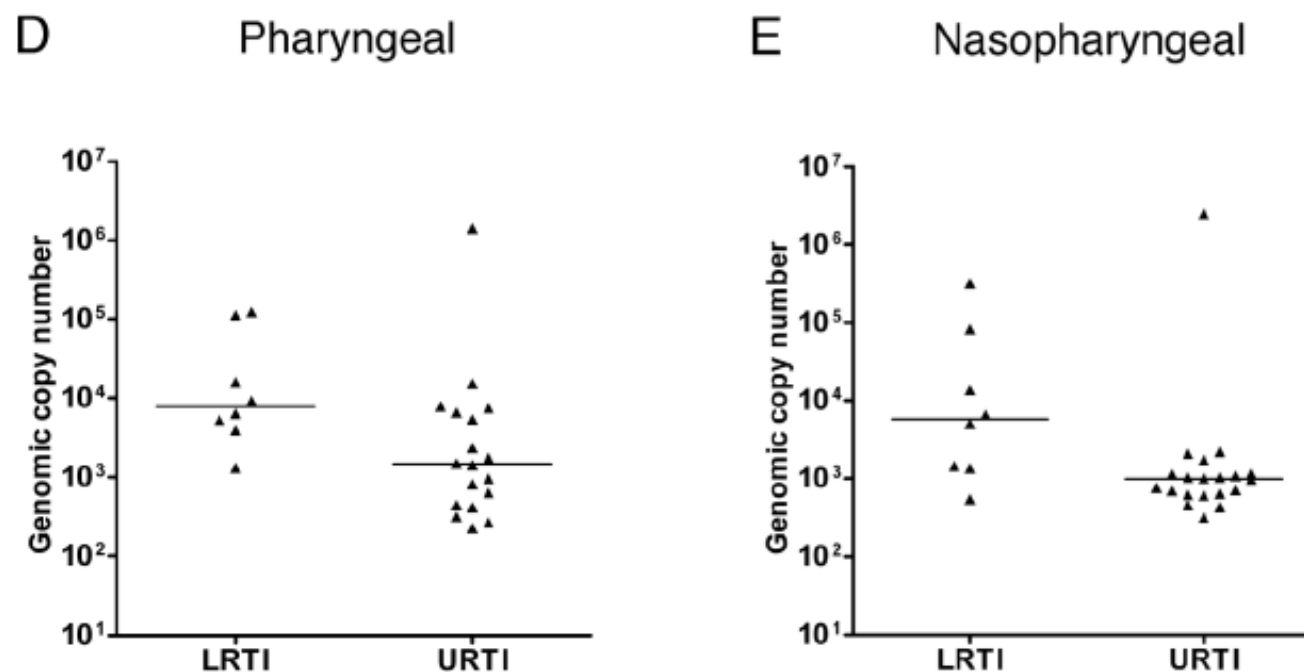
*Plos Med* 2013;10:e1001444



Pas de différence selon le type de prélèvement pour portage et infection

# Charges virales selon le type d'infection

*Plos Med 2013;10:e1001444*

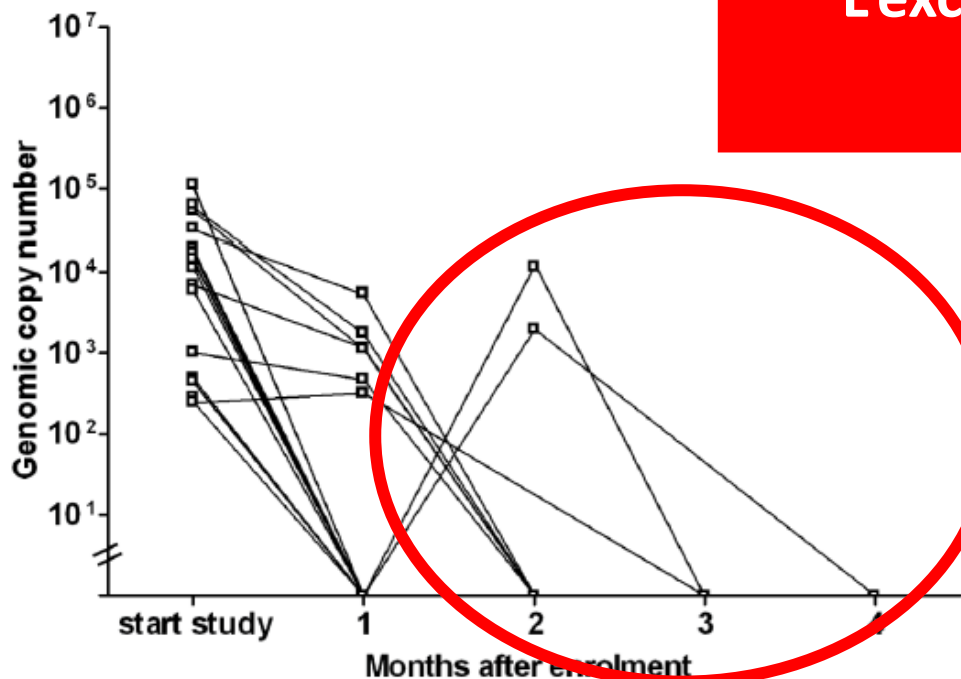


CV médiane + élevée pour infection VAI,  
mais distributions larges dans les deux types d'infections

# Durée de portage (étude ancillaire)

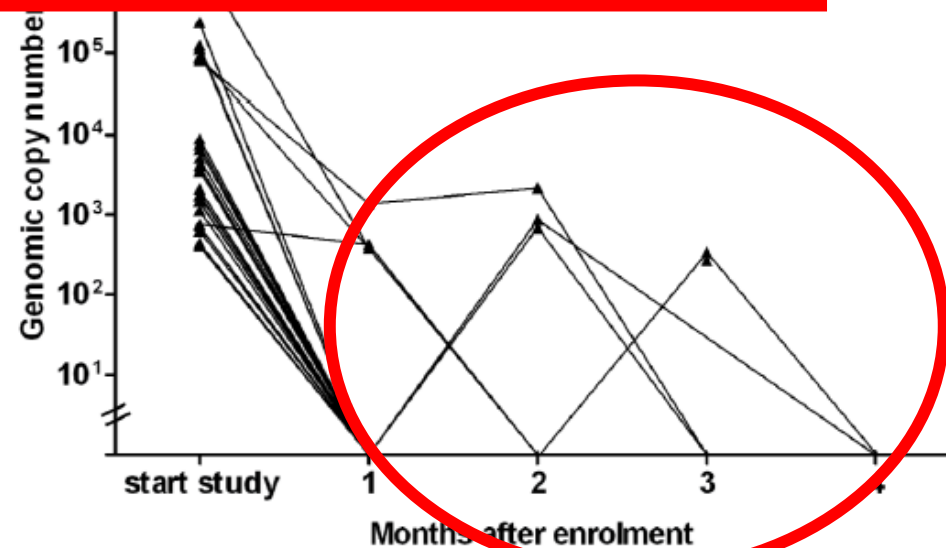
*Plos Med 2013;10:e1001444*

A Asymptomatic



L'excrétion peut être prolongée chez certains patients

71% négatifs à 1 mois

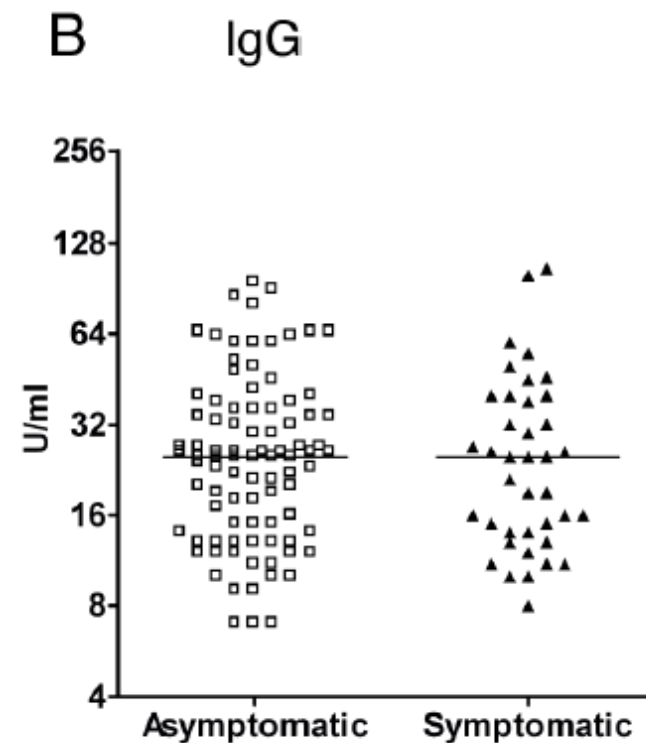
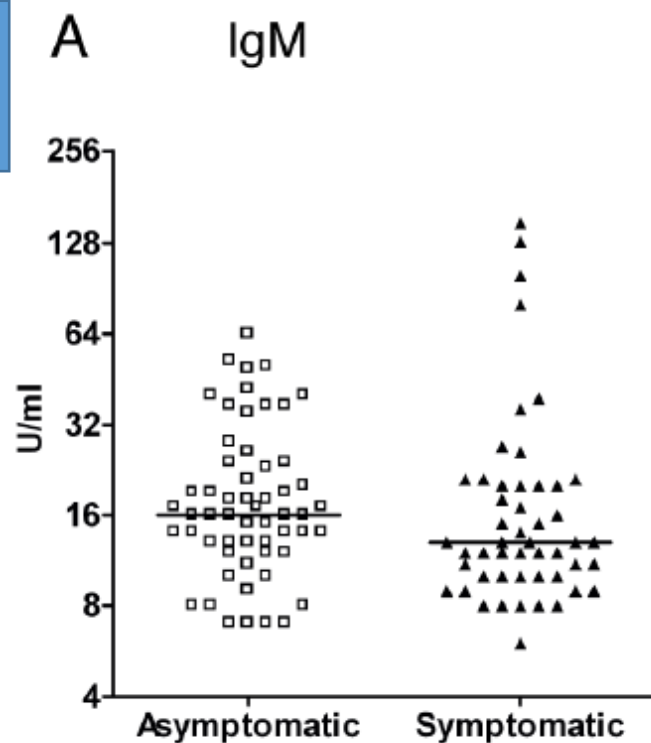


86% négatifs à 1 mois

# Portage vs infection : sérologie

*Plos Med 2013;10:e1001444*

Titre Ac



Positivité de la  
sérologie

**12,6% vs 9,2%**  
 **$P = 0,23$**

**25,1% vs 14,2%**  
 **$P < 0,001$**

# Portage versus infection : conclusions

- Mycoplasma pneumoniae est un **germe de portage**
- Très peu de corrélation entre PCR et sérologie
- Excrétion peut être prolongée



L'interprétation d'un test positif doit être précautionneuse !

# Rôle exact des germes que l'on trouve ?

**Table 6.** Virology results in 202 study participants.

| Virus             | Asymptomatic Participants ( <i>n</i> = 112) |                    | Symptomatic Participants ( <i>n</i> = 90) |                    | <i>p</i> -Value |
|-------------------|---|--------------------|---|--------------------|-----------------|
|                   | Ct Value, Median (IQR)                      | <i>n</i> (Percent) | Ct Value, Median (IQR)                    | <i>n</i> (Percent) |                 |
| Influenza A virus | 20.1  | 1 (0.9)            | 23.0 (18.0–33.8)                          | 6 (6.7)            | 0.03            |
| Coronavirus OC43  | 35.6 (28.7–37.3)                            | 5 (4.5)            | 29.0 (24.1–32.9)                          | 5 (5.6)            | 0.72            |
| Coronavirus NL63  | 37.5 (34.8–38.6)                            | 6 (5.4)            | 27.5 (27.0–38.0)                          | 7 (7.8)            | 0.49            |
| Bocavirus         | 30.9 (26.5–34.0)                            | 16 (14.3)          | 32.0 (28.8–34.0)                          | 3 (3.3)            | 0.008           |
| Adenovirus        | 31.9 (27.9–33.8)                            | 17 (15.2)          | 27.0 (26.2–30.5)                          | 9 (10.0)           | 0.28            |

Deux pathogènes ou plus chez 55,5% des symptomatiques  
... et 56% des asymptomatiques !

# Mycoplasme et atteinte respiratoire

# Infections respiratoires hautes

Portage ?

- Pharyngite non streptococcique : n = 184 enfants
- Réalisation PCR/sérologie mycoplasme :
- **24 %** PCR et/ou sérologie mycoplasme positive
- ATCD : > 3 épisodes de pharyngites dans les 6 mois :
  - 54,2% groupe mycoplasme +
  - 2,8% groupe mycoplasme -

*Eur J Clin Microbiol Infect Dis 2002;21:607-10*

Mais 3,8% dans autre étude espagnole

*Enferm Infecc Microbiol Clin 2009,27(7):403-5*

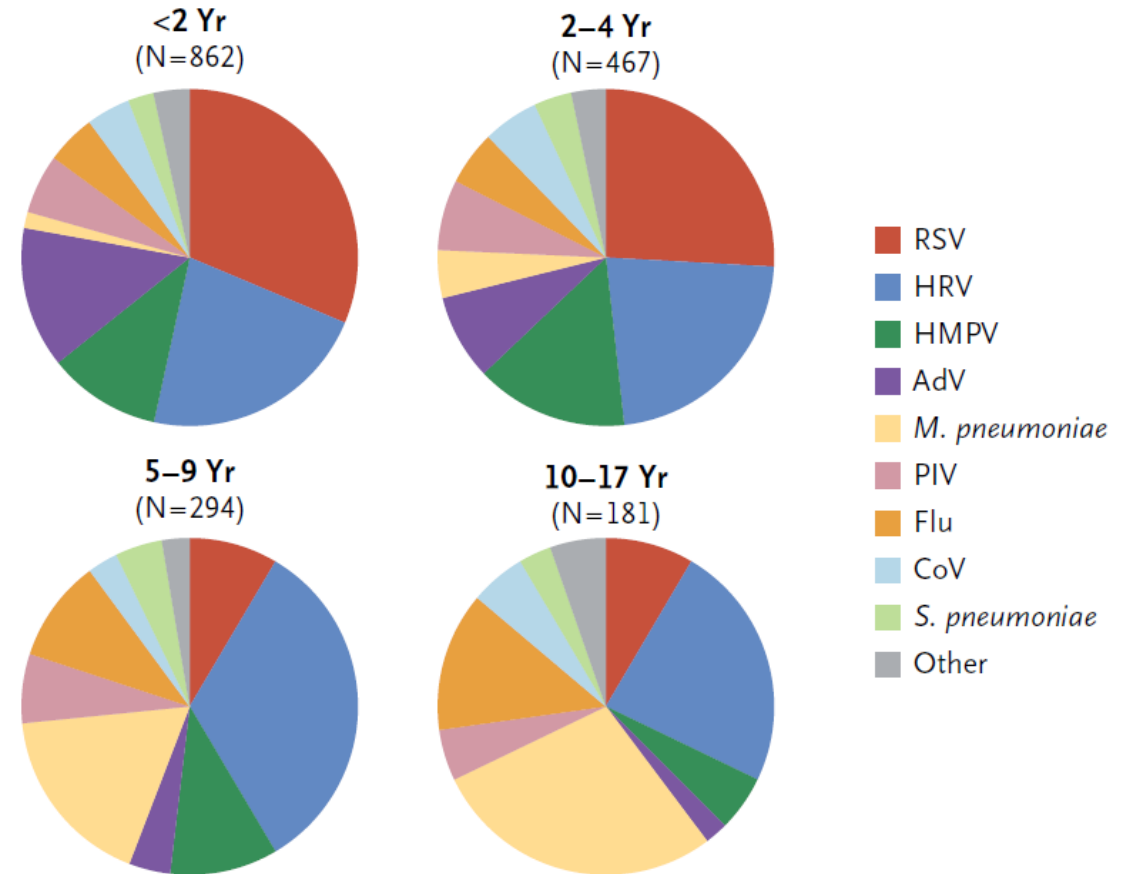
**Pour la pratique : !**

**Pas de recherche de mycoplasme devant une pharyngite même récidivante**

# *M. pneumoniae* et pathologie respiratoire

- 10-40% des infections respiratoires basses en pédiatrie
- 1<sup>ère</sup> bactérie des pneumopathies communautaires hospitalisées aux USA (TN, UT), surtout chez 10-17 ans
- Clinique : toux qui traîne...

C Detection According to Age Group



# Clinical symptoms and signs for the diagnosis of *Mycoplasma pneumoniae* in children and adolescents with community-acquired pneumonia (Review)

## AUTHORS' CONCLUSIONS

### Implications for practice

This review has found that *M. pneumoniae* cannot be reliably diagnosed in children and adolescents with community-acquired pneumonia based on the absence or presence of individual clinical symptoms and signs. Although the absence of wheeze is a statistically significant diagnostic indicator, it does not have sufficient diagnostic value to guide empirical macrolide antibiotic treatment

published. Clinicians should therefore consider other factors, including previous antibiotic use and population-based data on *M. pneumoniae* incidence, to help them estimate the likelihood of *M. pneumoniae* infection in the context of a clinical consultation.

# Quelles anomalies radiographiques ?



|   |   |  |  |
|---|---|--|--|
| Radiological images:<br>CT scan           | Bronchovascular bundles thickening  | Centrilobular nodules  | Consolidation,<br>Ground-glass attenuation                     |
| Pathological findings:<br>biopsy specimen | Bronchitis characterized by inflammatory cells and plasma cells in the large airway walls | Cellular bronchitis in the small airways with exudate in the lumen | Neutrophils and exudate within bronchiolar and alveolar lumens |
| Frequency (n= 91)                         | 75%   | 65%  | 66%  |

# *M. pneumoniae* chez < 1an



Chine : 217

**Table 6** Pathogens detected in respiratory tract co-infection.

| Parameters                | Pathogens   | Number | > 12 months<br>(n = 36) | P                  |
|---------------------------|---|--------|-------------------------|--------------------|
| Clinical symptoms         | <i>M. pneumoniae</i> + respiratory syncytial virus  | 5      |                         |                    |
| Cough (n [%])             | <i>M. pneumoniae</i> + adenovirus                   | 2      | (97.22%)                | 1.00 <sup>a</sup>  |
| Tachypnea (n [%])         | <i>M. pneumoniae</i> + parainfluenza virus 3        | 2      | 5.56%)                  | 0.089 <sup>a</sup> |
| Clinical signs            | <i>M. pneumoniae</i> + parainfluenza virus1         | 1      |                         |                    |
| Fever (>37.5 °C) (n [%])  | <i>M. pneumoniae</i> + influenza virus A + hBov     | 1      | (63.89%)                | 0.006              |
| Maximum temperature >     | <i>M. pneumoniae</i> + influenza virus A + hBov     | 1      | (41.67%)                | 0.033 <sup>a</sup> |
| Length of fever ≥3 days ( | <i>M. pneumoniae</i> + hBov                         | 2      | (44.44%)                | 0.013 <sup>a</sup> |
| Wheezing (n [%])          | <i>M. pneumoniae</i> + hMPV                         | 1      | (50%)                   | 0.49               |
| Gastrointestinal symptom  | <i>M. pneumoniae</i> + parainfluenza virus 2        | 1      | 25%)                    | 0.68               |
|                           | <i>M. pneumoniae</i> + parainfluenza virus 3 + hBov | 1      |                         |                    |
|                           | <i>M. pneumoniae</i> + hMPV + hBov                  | 1      | rate                    |                    |

# Mycoplasme et asthme

**Table 2. Clinical and spirometric data for children hospitalized for their first asthma attack.**

| Variable                                     | <i>M. pneumoniae</i> –<br>or <i>C. pneumoniae</i> –<br>positive patients<br>( <i>n</i> = 29) | <i>M. pneumoniae</i> –<br>or <i>C. pneumoniae</i> –<br>negative patients<br>( <i>n</i> = 22) |
|--|--|--|
| No. of male subjects/no. of female subjects  | 15/14  | 15/7   |
| Age, mean years                              | 5.7  | 5.5  |
| Familial asthma                              | 12 (41)  | 15 (68)  |
| Atopic dermatitis                            | 2 (7) <sup>a</sup>   | 12 (55) <sup>a</sup>   |
| Serum IgE of >200 IU/mL                      | 11 (30) <sup>a</sup>   | 15 (68) <sup>a</sup>   |
| Relapse of asthma ≤1 year after first attack | 18 (62) <sup>a</sup>   | 6 (27) <sup>a</sup>  |

# M. Pneumoniae et risque asthme

- Etude nationale
- 1591 infectés par an
- Score propensity

TABLE E3. Incidence of asthma after matching

Variables

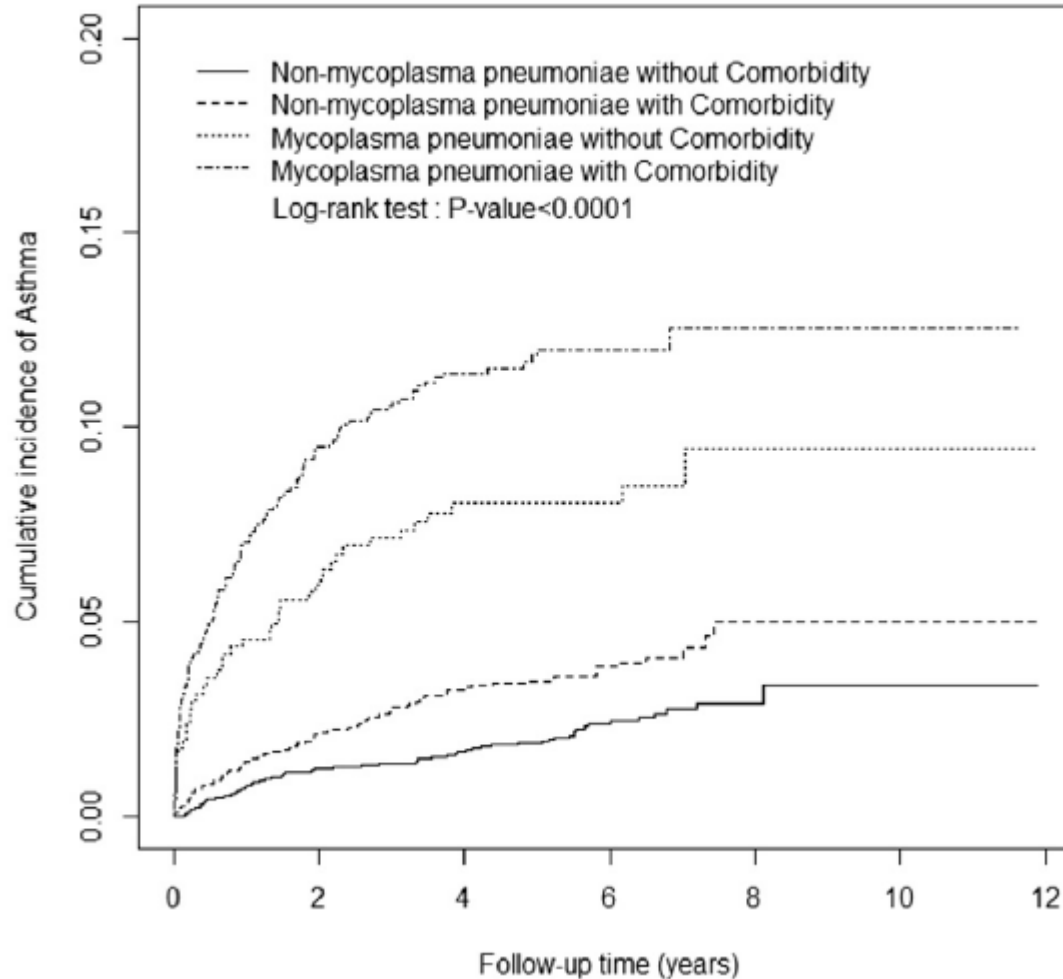
Overall asthma

Early-onset asthma

Late-onset asthma

Early-onset asthma, Age at diagnosis (years).

\*P < .001.



2000-2008)

infectés sur genre, âge,

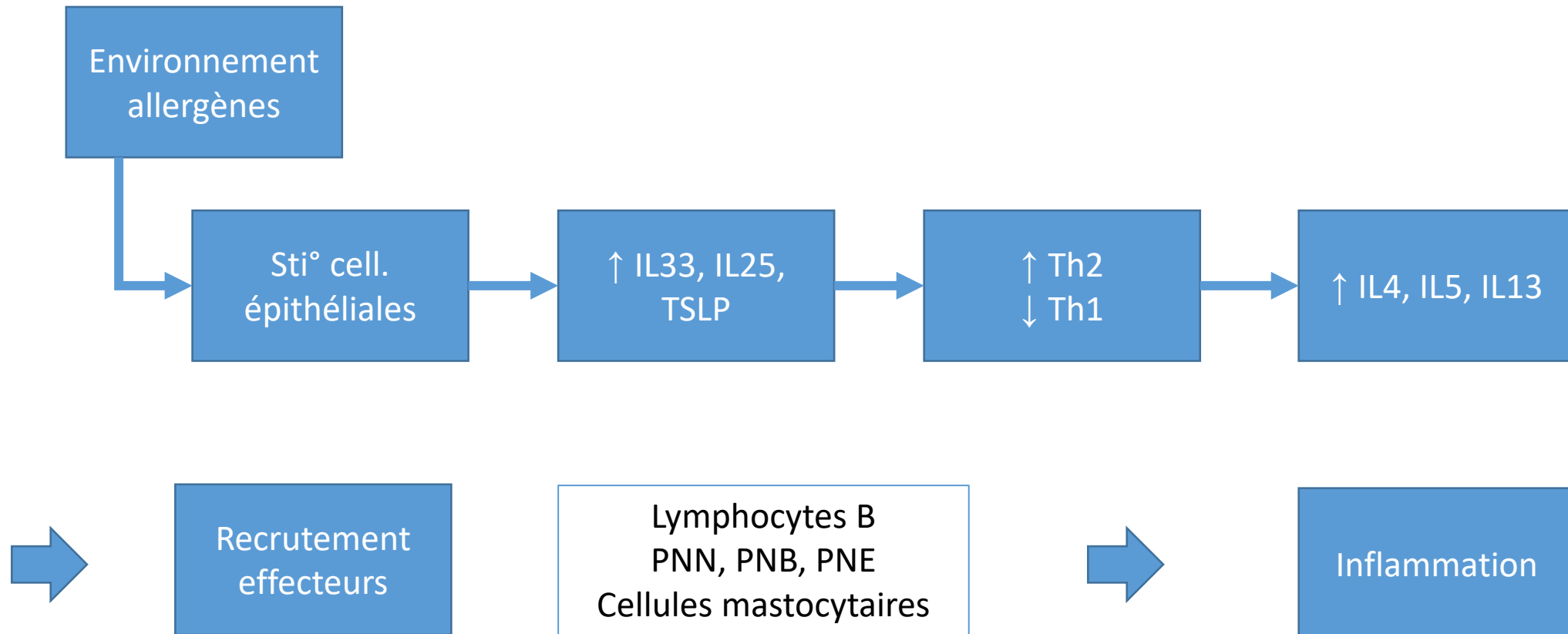
ou après âge de 12 ans)

*Mycoplasma pneumoniae* cohorts after propensity score

| Rate | Compared with non- <i>M pneumoniae</i> |                   |
|------|--|-------------------|
|      | IRE (95% CI)                           | HR (95% CI)       |
| .06  | 3.63 (3.07-4.29)*                      | 3.86 (2.94-5.09)* |
| .04  | 3.54 (2.96-4.24)*                      | 3.78 (2.66-5.38)* |
| .23  | 3.76 (3.15-4.49)*                      | 4.00 (2.57-6.20)* |

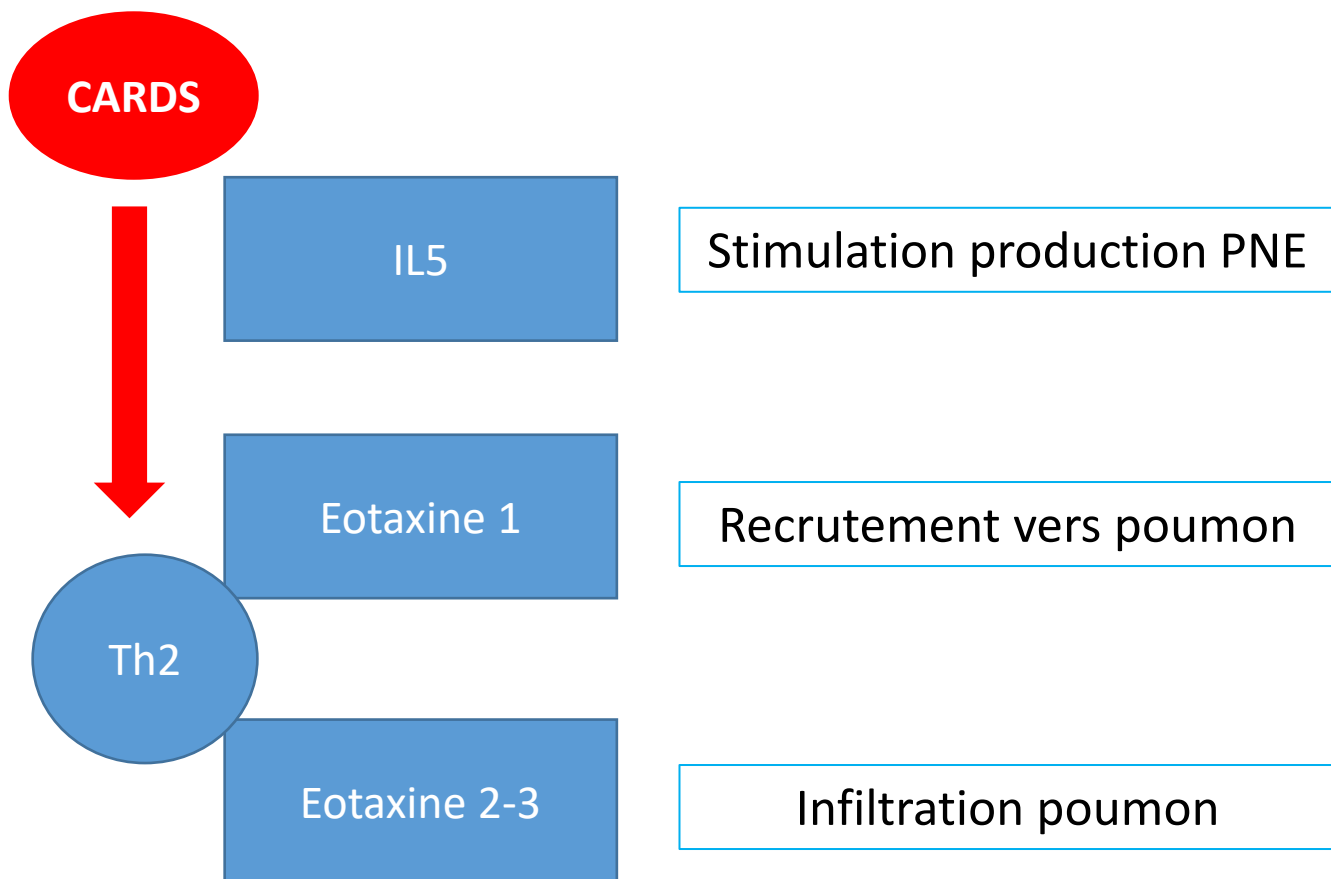
Rate, incidence rate (per 1000 person-years); \*P < .001.

# *M. pneumoniae* et exacerbation d'asthme



# Les PNE dans l'asthme

*J Clin Cell Immunol 2018;9(4)*



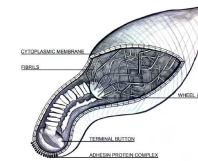
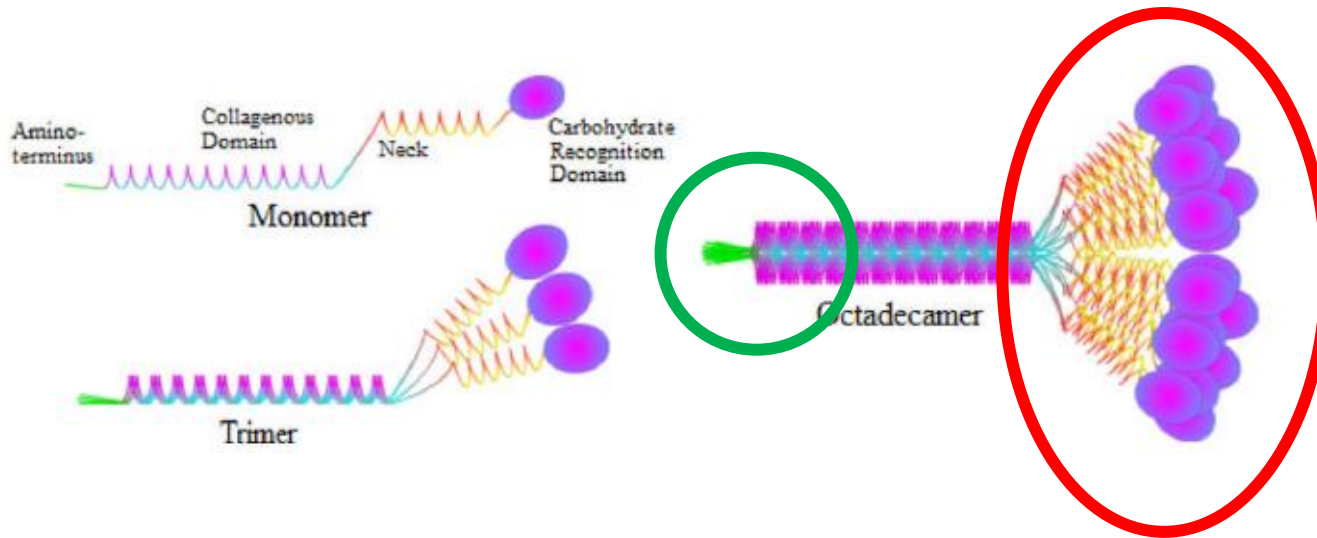
- Granules cytotoxiques
  - Major basic protein (MBP)
  - Eosinophil cationic protein (ECP)
  - Eosinophil peroxydase (EPO)
  - Eosinophil-derived neurotoxin (EDN)

Dégranulation mastocytes, PNB

- Cellules présentatrice Ag

Régulation Th1 / Th 2

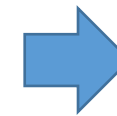
# Protéine A du surfactant pulmonaire (SP-A)



↓ Production EPO par PNE

Stimule les macrophages alvéolaires

Sert de ligand aux micro-organismes, allergènes et cellules apoptotiques

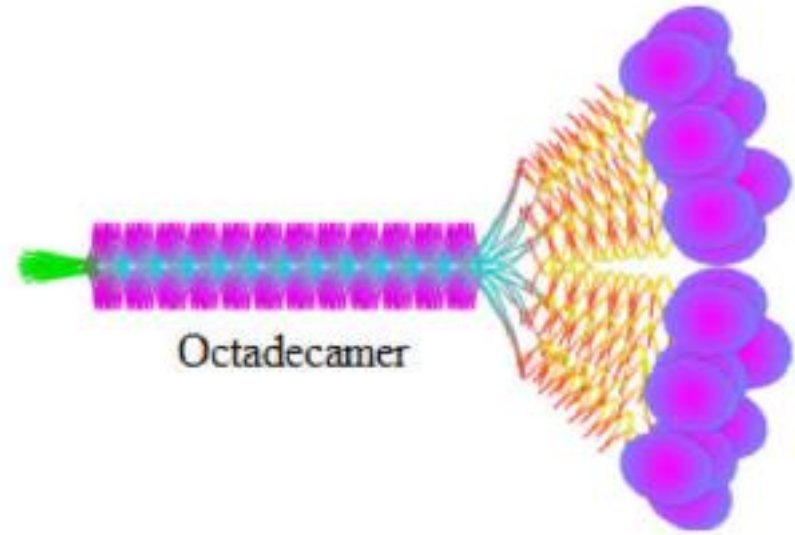


Moins d'inflammation

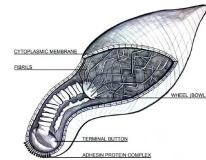
# Mutation SP-A et M. pneumoniae

Mutation Q223 / K223  
Dans gène SP-A2

Gènes SP-A1  
et SP-A2



Balance pro-inflammatoire  
= Asthme



~~↓ Production EPO par PNE~~

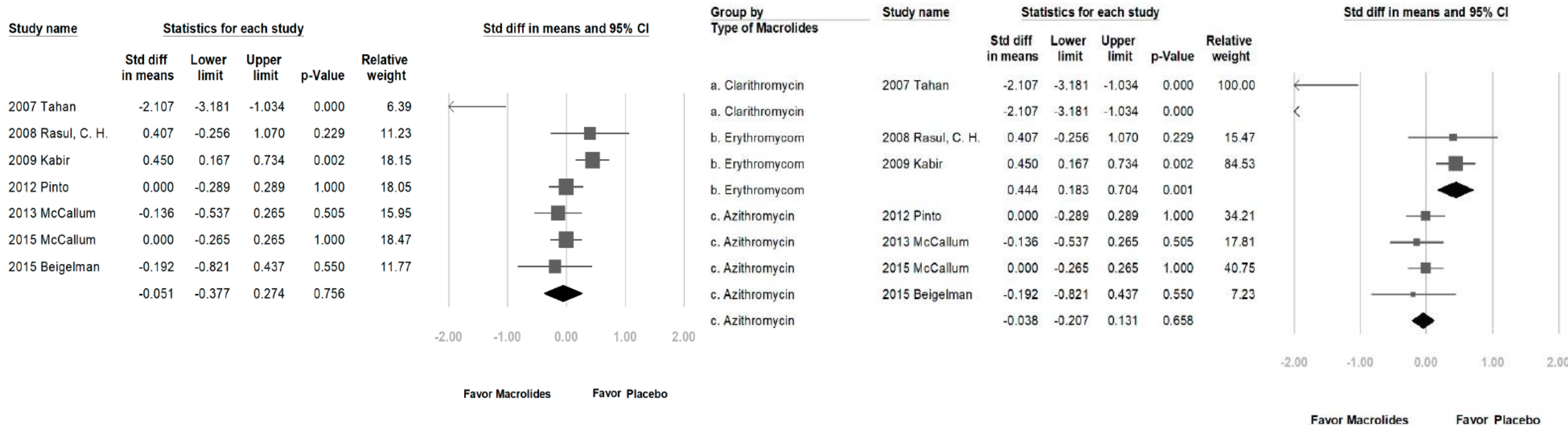
# Une place pour les macrolides devant wheezing ?

- Une **action sur mycoplasme** (retrouvé associé au wheezing)
- Une **action anti-inflammatoire**
  - ↓ PNN, IL8, IL6, IL1, TNF, ECP
  - Action sur balance Th1/2 : ↓ IL4, IL5, IL6



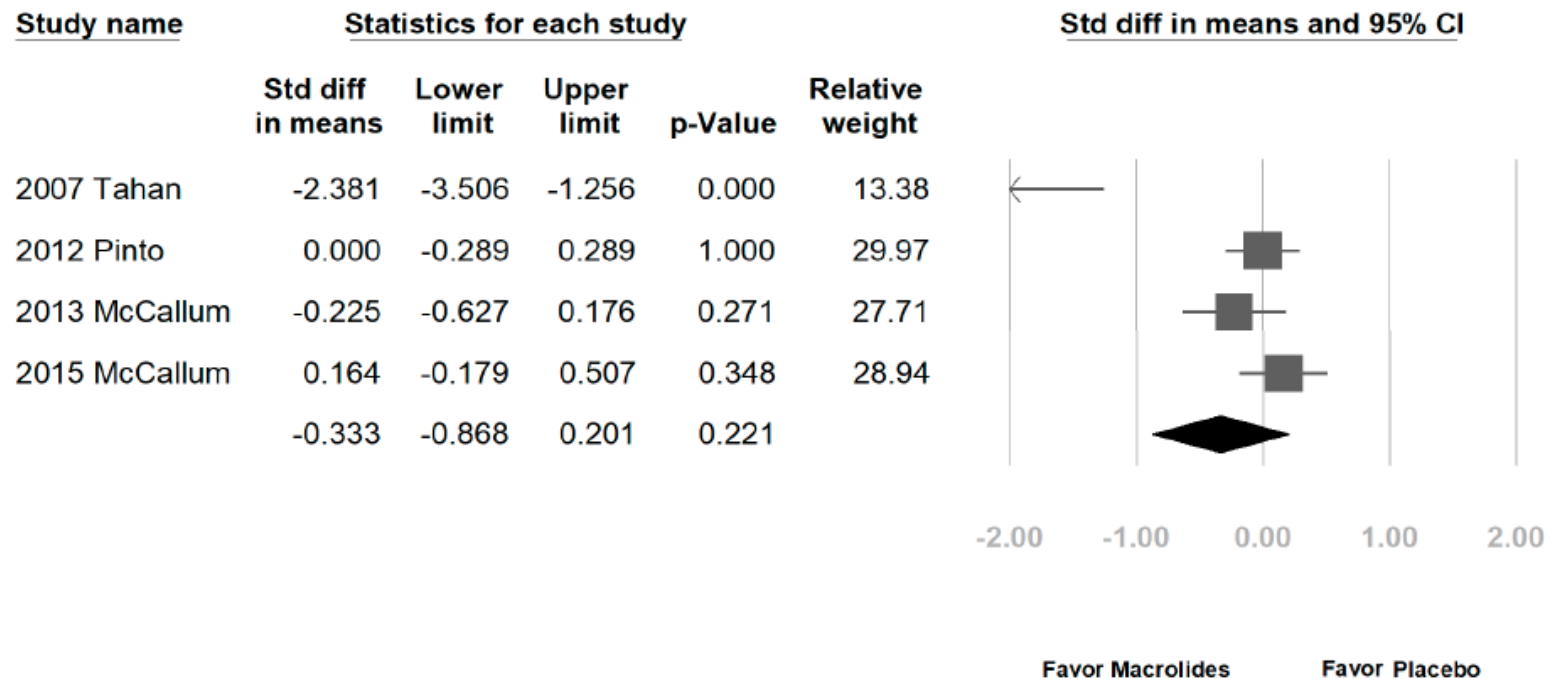
Utilité en systématique ?

# Une place pour les macrolides devant wheezing ?



**Aucun bénéfice sur durée d'hospitalisation**

# Une place pour les macrolides devant wheezing ?

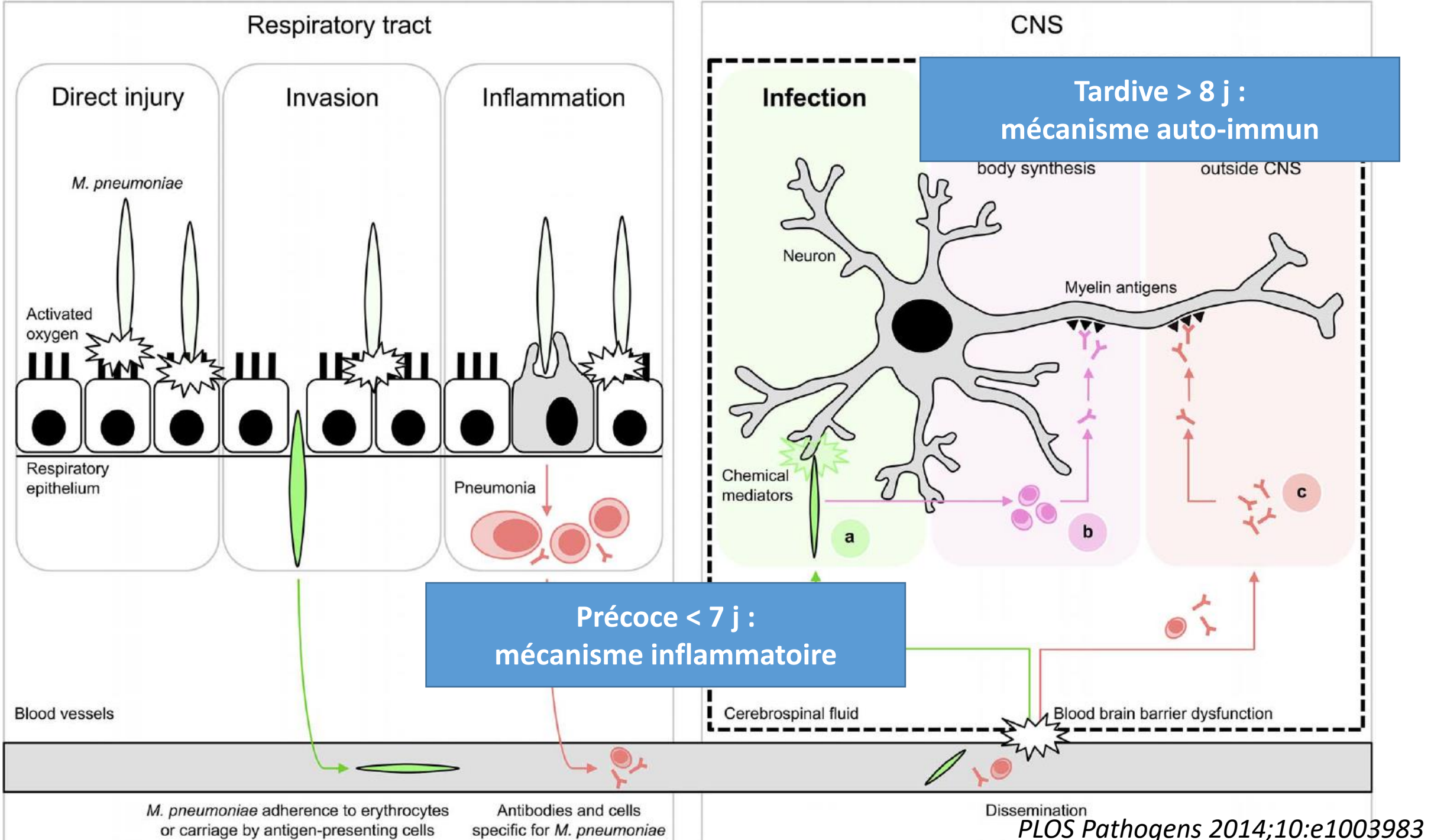


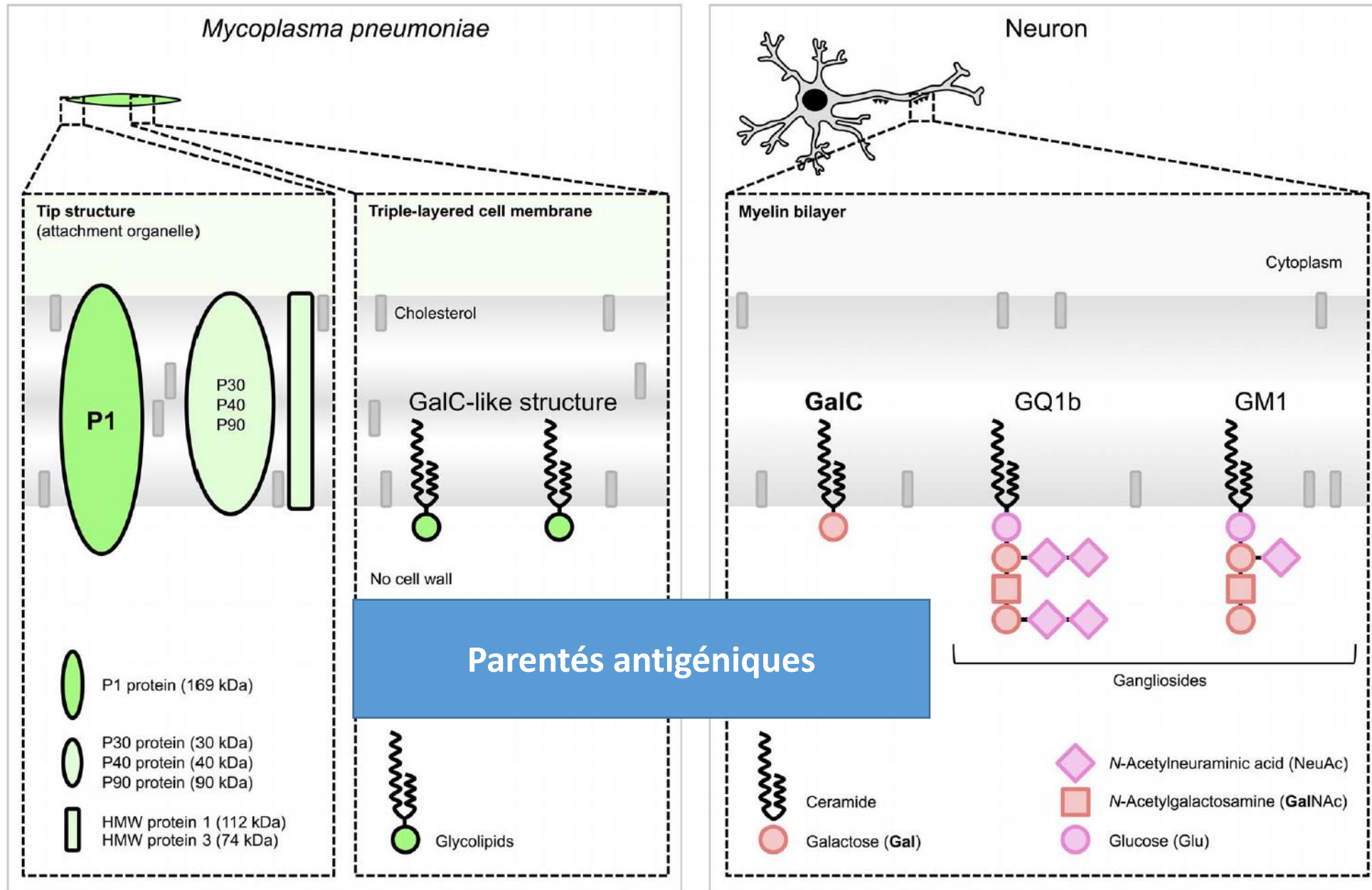
Aucun bénéfice sur durée de besoin en oxygène

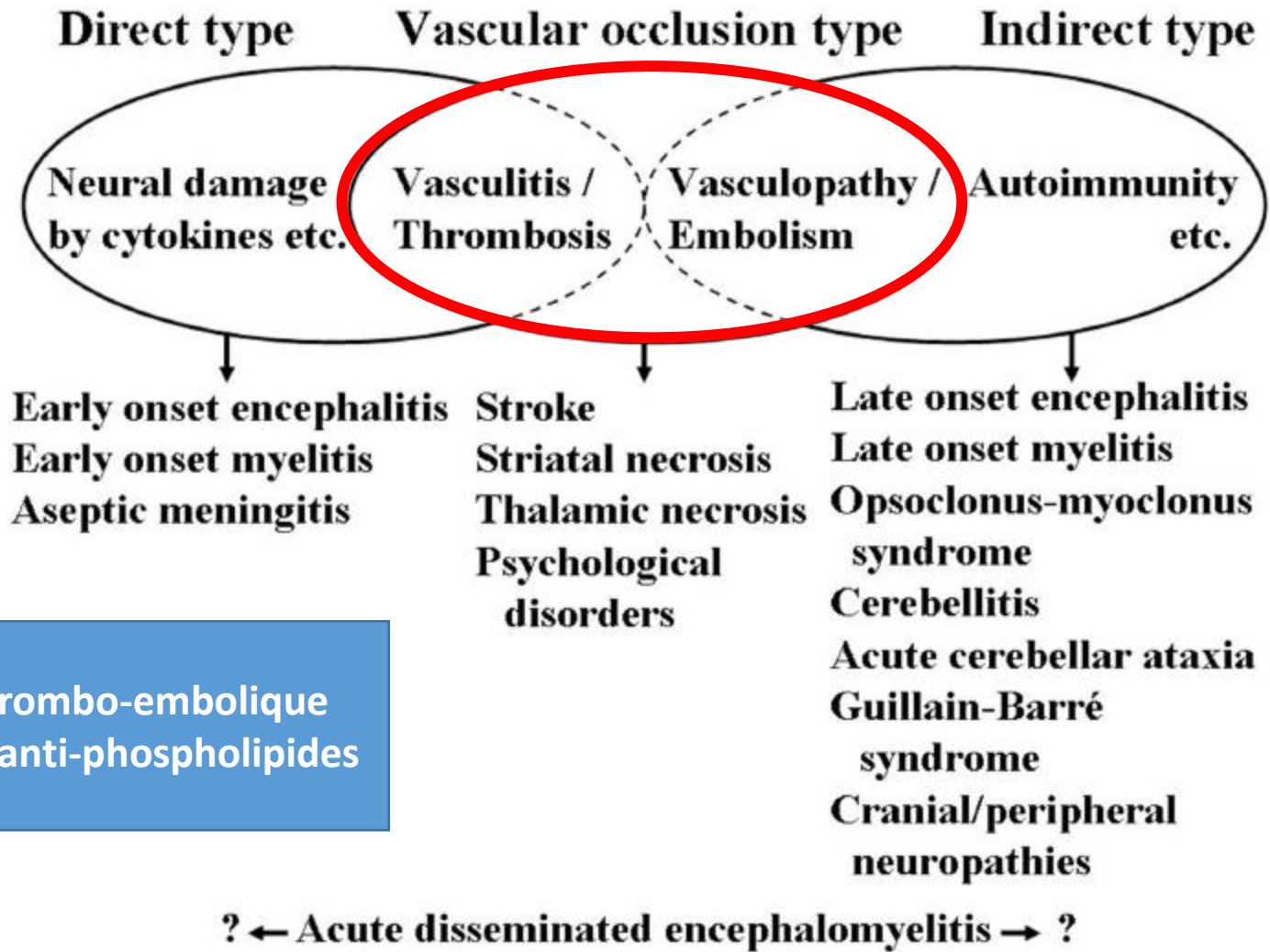
# Mycoplasme et atteinte neurologique

# *M. pneumoniae* et atteinte neurologique

- 5-10% des patients avec atteinte fébrile du SNC sont Mp+
- Encéphalite ++
- Méningite aseptique, myélite transverse, Guillain-Barré, AVC...
- Sérologie positive dans LCR (74% encéphalites à Mp), voire PCR+







Atteinte vasculaire, thrombo-embolique  
 Complexes immuns, Ac anti-phospholipides

# Mycoplasme et atteinte cutanéomuqueuse

# Atteintes cutanéomuqueuses

- 1-5% des infections à Mp
- Exanthème dans 17% des pneumonies à Mp
- Du rash... au Steven-Johnson
- « *Mycoplasma pneumoniae* – associated mucositis » (MPAM) ou syndrome de Fuchs

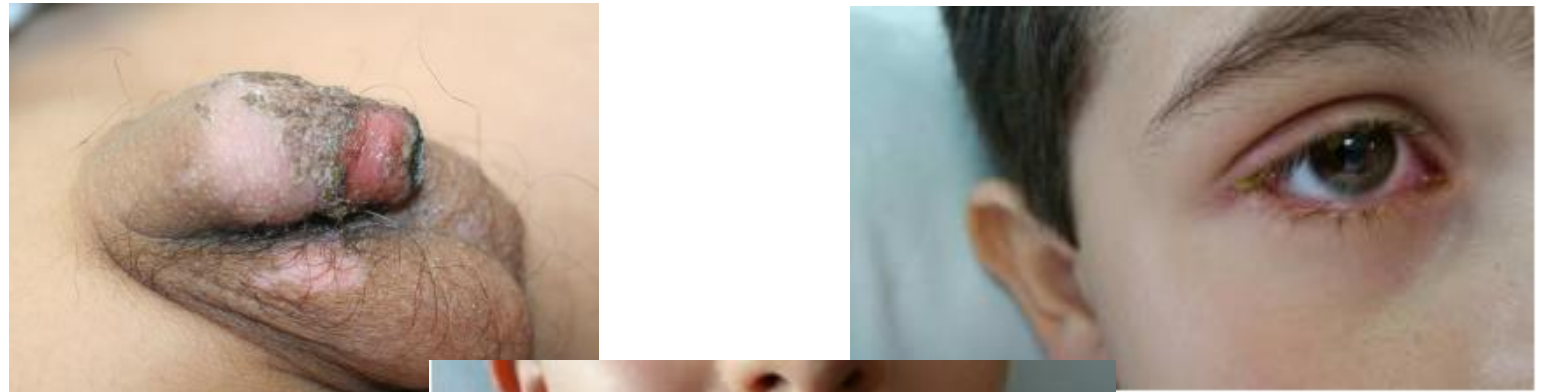
|             | Fuchs' syndrome | Varicella-like eruption | Vasculitis | Erythema nodosum |
|-------------|-----------------|-------------------------|------------|------------------|
| Patients, n | 37              | 8                       | 21         | 11               |
| Age, years  |                 |                         |            |                  |
| Childhood   | 21              | 5                       | 14         | 8                |
| Adulthood   | 16              | 3                       | 7          | 3                |
| Gender      |                 |                         |            |                  |
| Male        | 32 <sup>a</sup> | 6 <sup>a</sup>          | 12         | 2 <sup>b</sup>   |
| Female      | 5 <sup>a</sup>  | 2 <sup>a</sup>          | 9          | 7 <sup>b</sup>   |

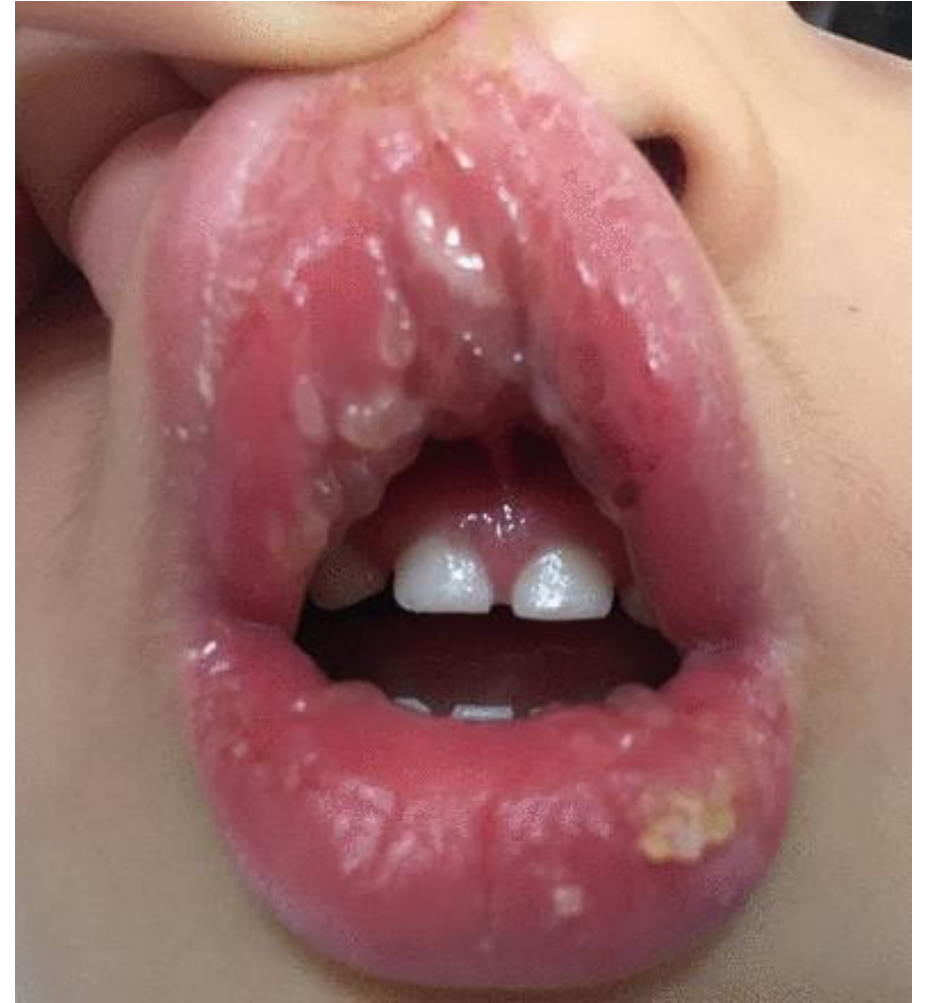
*Dermatology* 2015;231:152-7

**Table I.** Clinical characteristics of *Mycoplasma*-induced rash and mucositis

|                                    |            |
|------------------------------------|------------|
| Age, y                             | 11.9 ± 8.8 |
| Gender                             |            |
| Male                               | 121 (66)   |
| Female                             | 62 (34)    |
| Unknown                            | 19 (10)    |
| Morphology*                        |            |
| Vesiculobullous                    | 72 (77)    |
| Targetoid or atypical targetoid    | 45 (48)    |
| Papules                            | 13 (14)    |
| Macules                            | 11 (12)    |
| Meibomian                          | 9 (9)      |
| Cutaneous involvement              |            |
| Sparse                             | 61 (47)    |
| Absent                             | 42 (34)    |
| Moderate <sup>†</sup>              | 24 (19)    |
| Cutaneous distribution             |            |
| Acral                              | 42 (46)    |
| Generalized                        | 29 (31)    |
| Truncal                            | 21 (23)    |
| Mucosal involvement                |            |
| Oral                               | 168 (94)   |
| Ocular                             | 147 (82)   |
| Urogenital                         | 112 (63)   |
| No mucositis                       | 4 (2)      |
| Clinical outcomes                  |            |
| Full recovery                      | 104 (81)   |
| PIPA                               | 7 (6)      |
| Mucosal complications <sup>‡</sup> | 13 (10)    |
| Recurrence                         | 10 (8)     |
| Death                              | 4 (3)      |

*J Am Acad Dermatol 2015;72:239-45 et Clin Case Rep 2018;6(3):551-2*





*Pediatrics 2015;136:e386*

*BMJ Case Reports 2018;  
doi:10.1136/bcr-2017-223321*

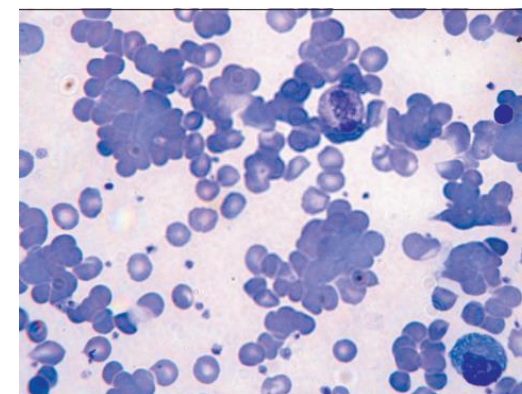
# Autres atteintes liées au Mycoplasme

# Anémie hémolytique à agglutinines froides

- Présence agglutinines froides : 50-60% des infections Mp
- **Précoces** (1 semaine jusqu'à 2-6 semaines)
- Le plus souvent anémie **pauci-symptomatique** (réticulocytose)
- Mécanisme direct (lyse paroi des GR) ou indirect (agglutination)
- Médié par le **complément**

- **Spécificité anti-I**

- Présent sur GR
- Présent sur sialo-oligosaccharides des cellules bronchiques (récepteur Mp)



**TABLE 1 |** Extrapulmonary manifestations due to *M. pneumoniae* infection classified according to the involved pathomechanisms.

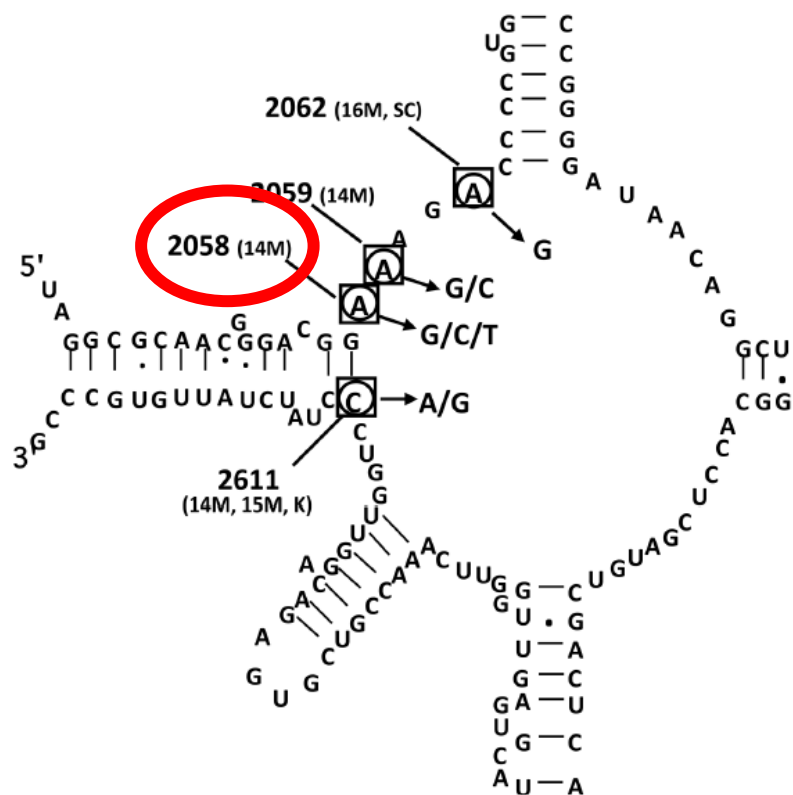
| Manifestations                         | Direct type <sup>a</sup>   | Indirect type <sup>b</sup>  | Vascular occlusion type <sup>c</sup>   | Undetermined <sup>d</sup>            |
|--|--|---|--|--------------------------------------|
| Cardiovascular system                  | Pericarditis, Endocarditis   | Myocarditis, Kawasaki disease   | <u>Cardiac thrombus</u> ,<br><u>Aortic thrombus</u>                          |                                      |
| Dermatological                         |  | Erythema multiforme, Urticaria, Anaphylactoid purpura, <u>EN</u> , <u>CLV</u> , <u>SJS</u> , <u>MPAM</u> , <u>SPD</u>   |  |                                      |
| Digestive organ                        | Early onset hepatitis  | Late onset hepatitis  | Pancreatitis   |                                      |
| Hematological/<br>Hematopoietic system |  | Autoimmune hemolytic anemia, Hemophagocytic syndrome, Thrombocytopenic purpura, Infectious mononucleosis  | Disseminated intravascular coagulation, Splenic infarct                      |                                      |
| Musculoskeletal system                 | Arthritis  |   |  | Rhabdomyolysis                       |
| Neurological                           | Early onset encephalitis, Early onset myelitis, Aseptic meningitis | Late onset encephalitis, Late onset myelitis, Guillain-Barré syndrome, Cranial/peripheral neuropathies, Cerebellitis, <u>Acute cerebellar ataxia</u> , <u>Opsoclonus-myoclonus syndrome</u> | Stroke, Psychological disorders, Striatal necrosis, <u>Thalamic necrosis</u> | Acute disseminated encephalomyelitis |
| Respiratory system                     |  |   | <u>Pulmonary embolism</u>  |                                      |
| Sensory organ                          | Otitis media   | Conjunctivitis, Iritis, Uveitis   | Sudden hearing loss  |                                      |
| Urogenital tract                       |  | Glomerulonephritis, IgA nephropathy   | Priapism,<br><u>Renal artery embolism</u>                                    |                                      |

# Le traitement

# Traitement ATB infections à *M.pneumoniae*

- Pas de paroi : R<sup>ce</sup> beta-lactamines, glycopeptides, fosfomycine
- Résistance naturelle : sulfamides, rifampicine, linezolide, polymyxine
- Sensibilité
  - MLSK ++
  - Tétracycline
  - FQ surtout moxifloxacine (Izilox<sup>®</sup>) et lévofloxacine (Tavanic<sup>®</sup>)

# Résistance macrolides



- Résistance par **mutation 23S rARN** et protéines ribosomales L4 et L22
- **A2058G et A2059G** : haut niveau résistance aux macrolides en C14 (clarithro) et 15 (azithro) (C16 (josa) aussi pour A2059G)
- **Pas de résistance croisée** avec tétracyclines et FQ
- Expression clinique : **persistance des signes**
- Options : doxycycline ou minocycline, FQ, pristinamycine?

# Messages

- *Mycoplasma pneumoniae* est un germe de **portage**
- **Sérologie et PCR** performantes... Mais à interpréter avec précaution
- Physiopathologie : **toxicité** directe, indirecte ou par **Ac auto-immuns**
- Rôle de **protéine P1** et de toxine **CARDS**
- **Grande diversité clinique ++**