

Perspectives des autorités de santé et modèle à compartiments

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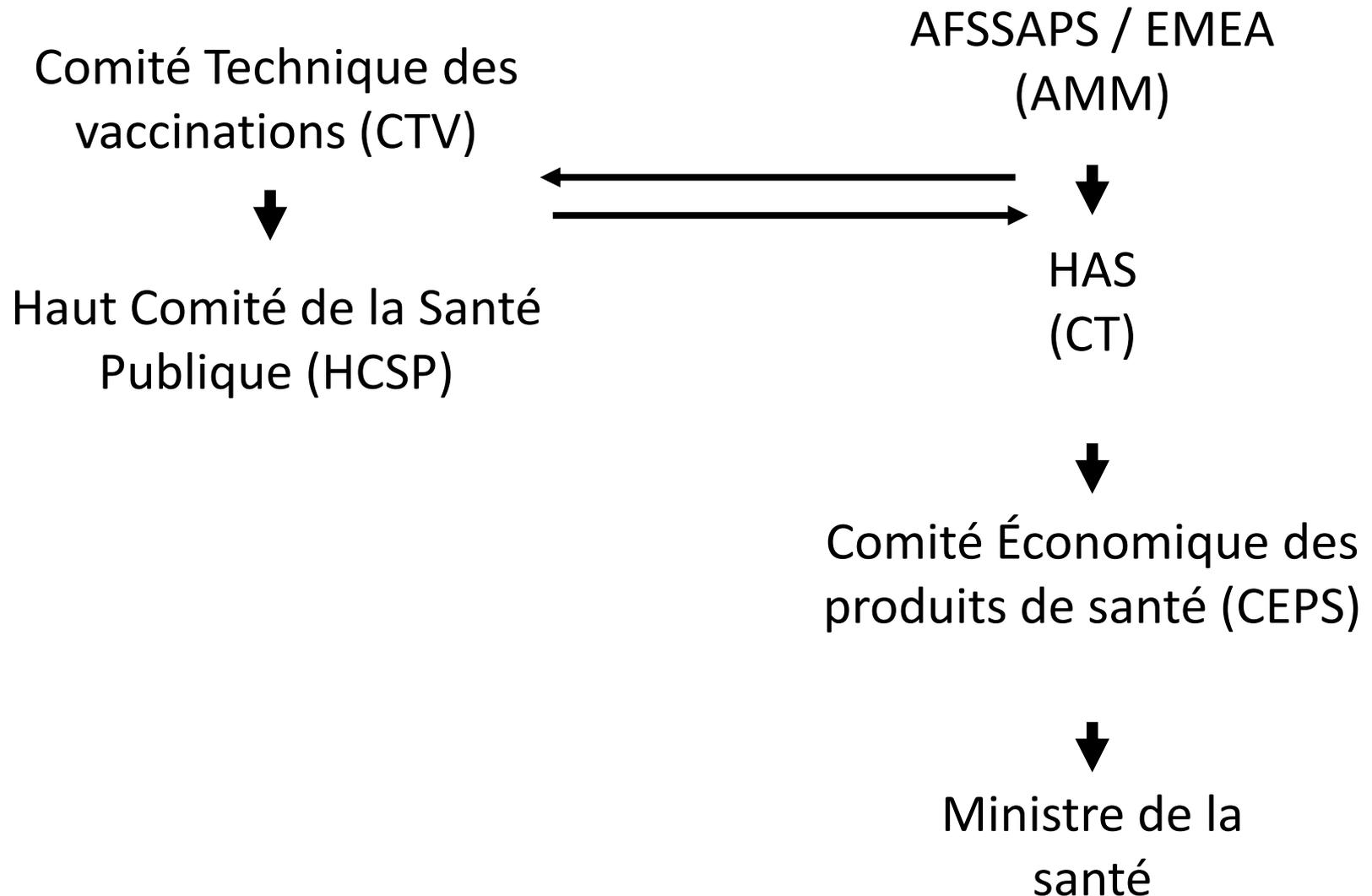
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Les spécificités du vaccin (1)

- Un contexte décisionnel particulier :
 - Des instances d'évaluation spécifiques (CTV, HCSP, à l'instar de nombreux pays)
 - Un statut particulier du vaccin ?
 - Des compétences particulières ?
 - Focus sur la stratégie plutôt que sur le produit
 - L'évaluation économique reconnue comme critère de décision [Cf. texte définissant les missions du CTV]
 - Les premières évaluations publiées concernaient la vaccination
 - Rapport coût / efficacité des vaccins (faible coût des vaccins, maladies très sévères, peu d'alternatives)
 - Un contexte qui se modifie [au regard de la population cible, du coût des vaccins, du fardeau des maladies, des stratégies alternatives]

L'accès au marché



Les spécificités du vaccin (2)

- L'existence d'externalités :
 - La vaccination a des effets directs et indirects positifs [réduction de la circulation du pathogène au sein de la population] ou négatifs [déplacement en âge des cas, remplacement sérotypique]
 - Ces effets doivent être pris en compte dans l'évaluation afin de ne pas sous- (ou sur-) estimer les bénéfices associés à la vaccination

EXTERNALITES → CHOIX DU MODELE

Les spécificités du vaccin (3)

- Les effets de long terme :
 - La vaccination produit des effets de long terme (retour à l'équilibre du système). Le choix de l'horizon temporel est déterminant dans le processus d'évaluation [effets de lune de miel]
 - Quelle pertinence du point de vue du décideur ?
 - Prise en compte de l'incertitude ?
 - Quelle est la valeur présente des bénéfices et des coûts survenant à long terme ? [Actualisation]

EFFETS DE LONG TERME →

**HORIZON TEMPOREL
INCERTITUDE
ACTUALISATION**

La place de la modélisation dans l'évaluation dans l'évaluation des stratégies vaccinales

Qu'est-ce qu'un modèle ?

- Un modèle = représentation mathématique simplifiée de la réalité qui permet d'aller au-delà des données fournies par les essais cliniques
- Un modèle → arbitrage entre réalisme et complexité (sous la contraintes des connaissances du moment et des données disponibles)
- Le modèle permet de mettre en cohérence les données disponibles, de nature diverse (clinique, épidémiologique, sur l'observance et/ou la persistance, économique...)
- Le modèle permet de quantifier l'incertitude

Les différents types de modèles

Statique

Dynamique

Ouvert

Fermé

Déterministe

Stochastique

Agrégé

Individu-centré

Temps discret

Temps continu

Le choix du modèle

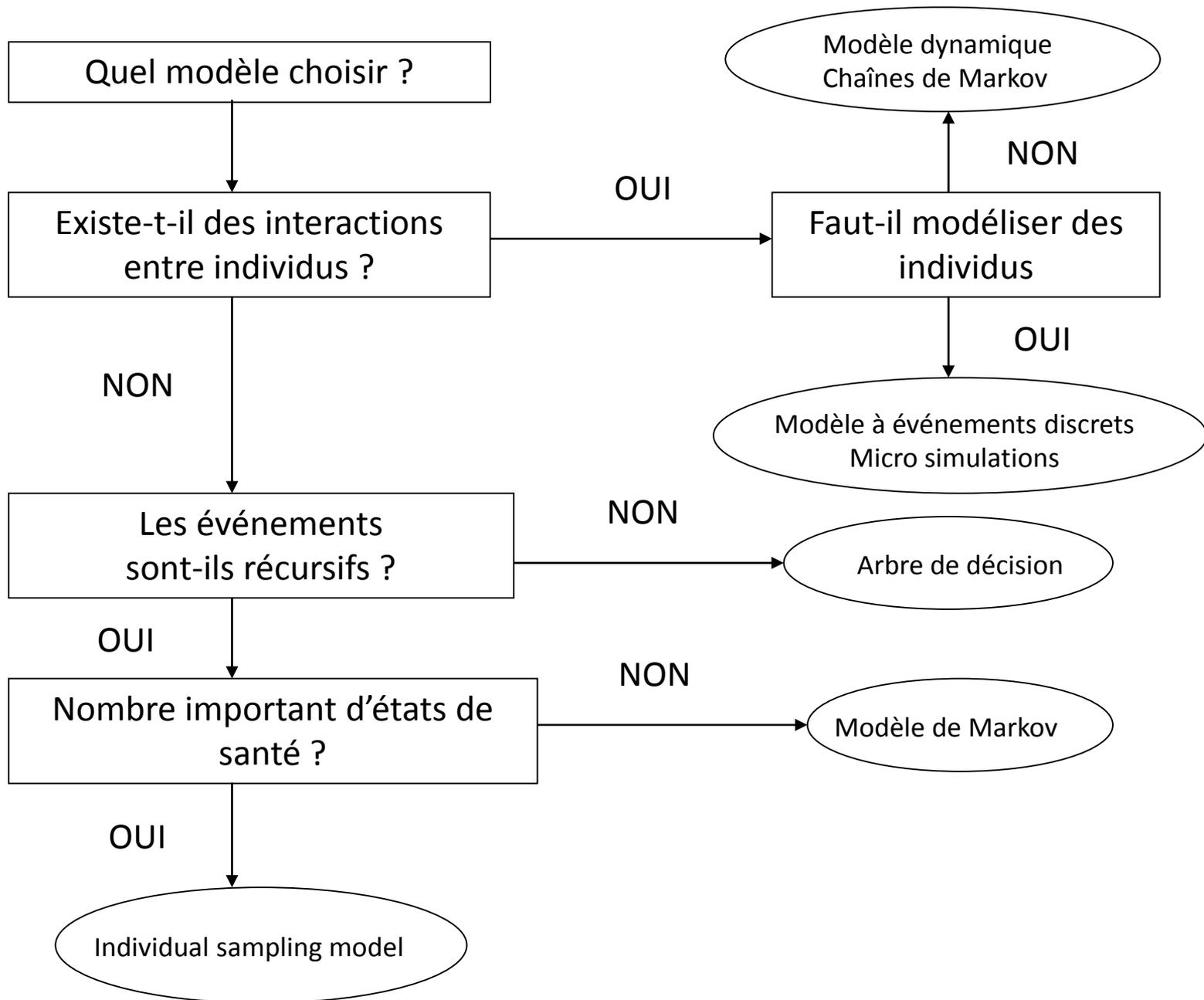
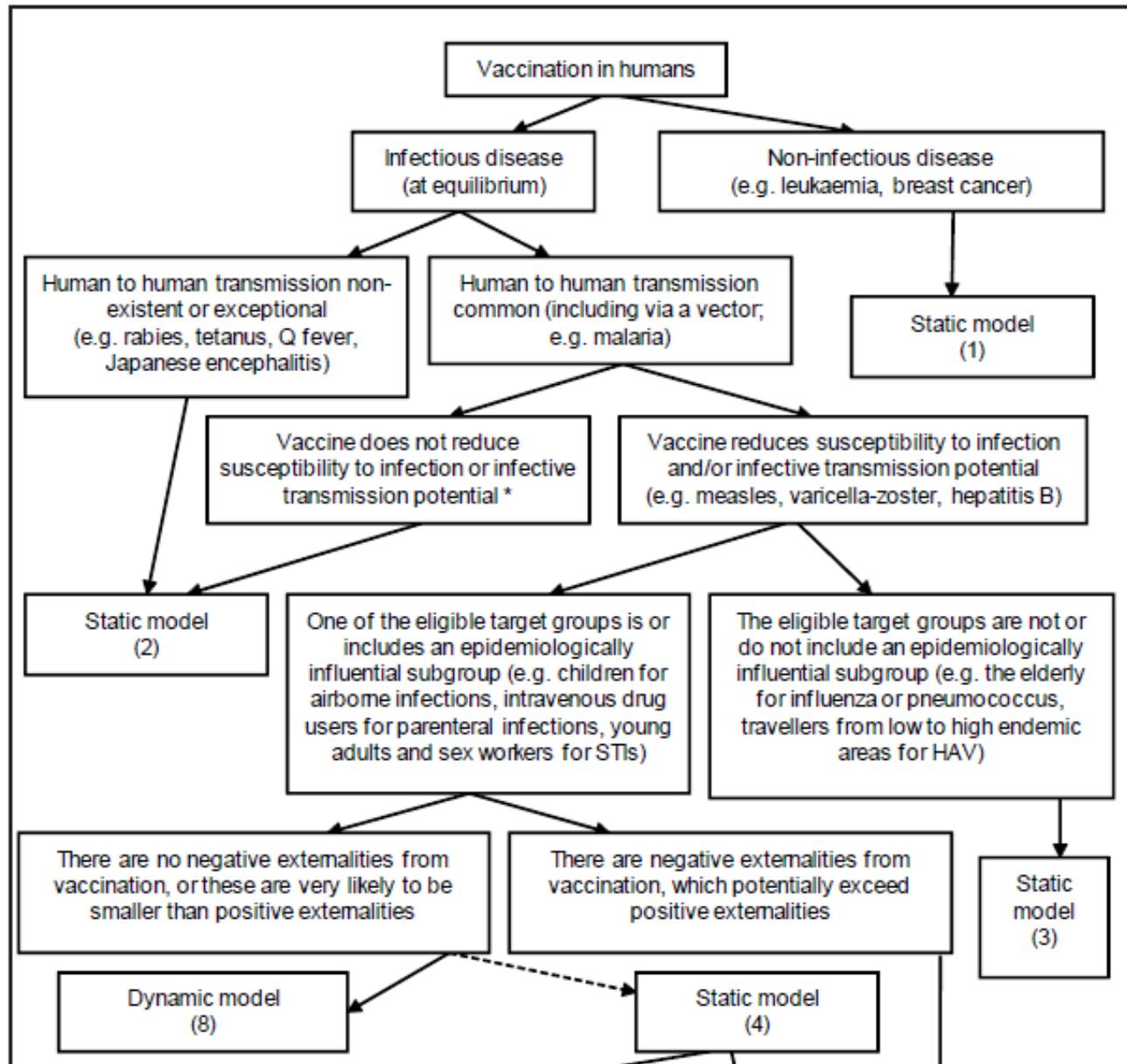
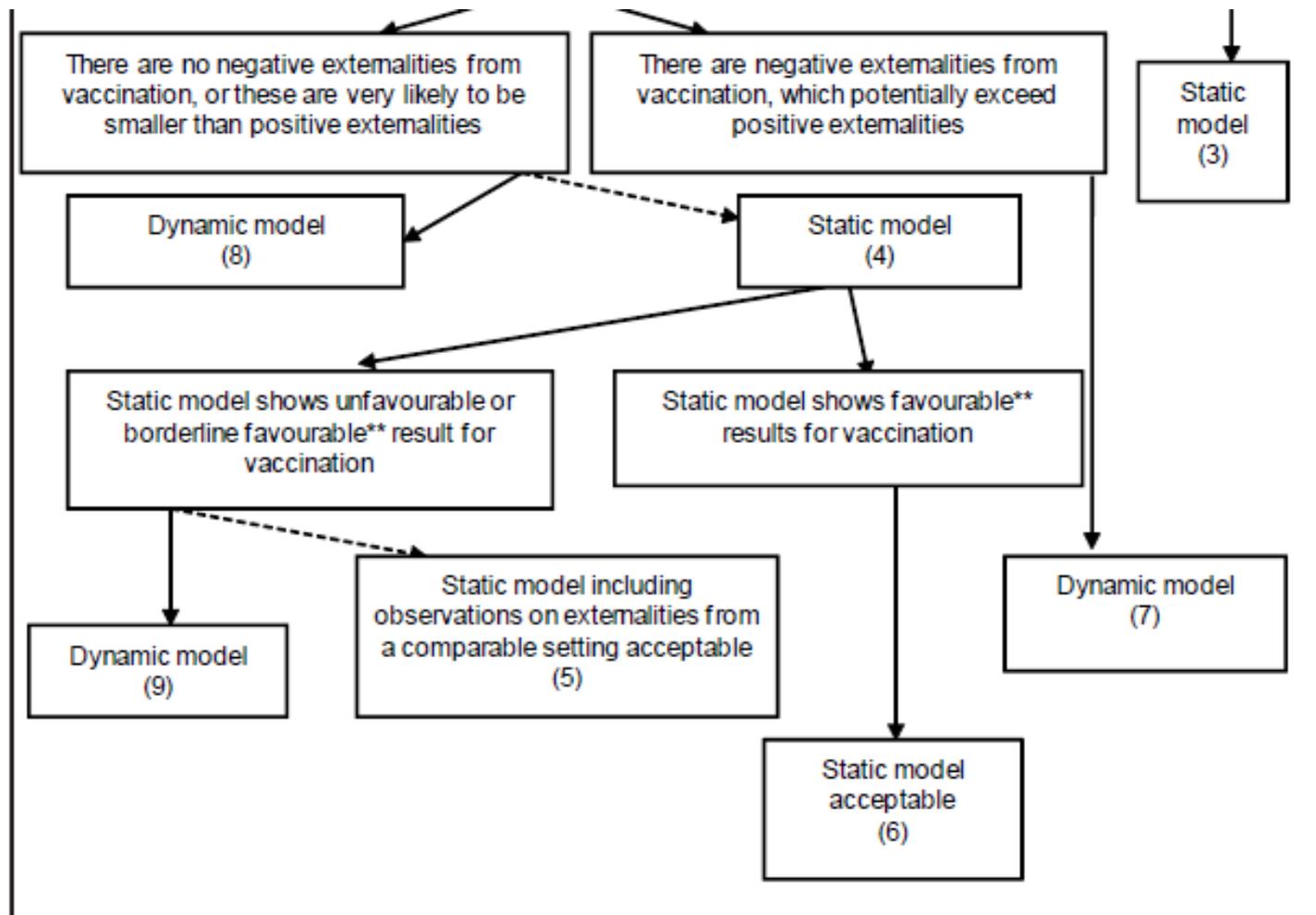
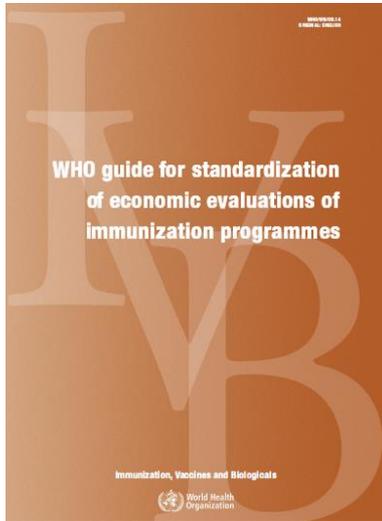


Figure 3: Flow chart to help determine when dynamic or static models are appropriate





Recommendations OMS (2008)



6.5 Recommendations

The mathematical model should be:

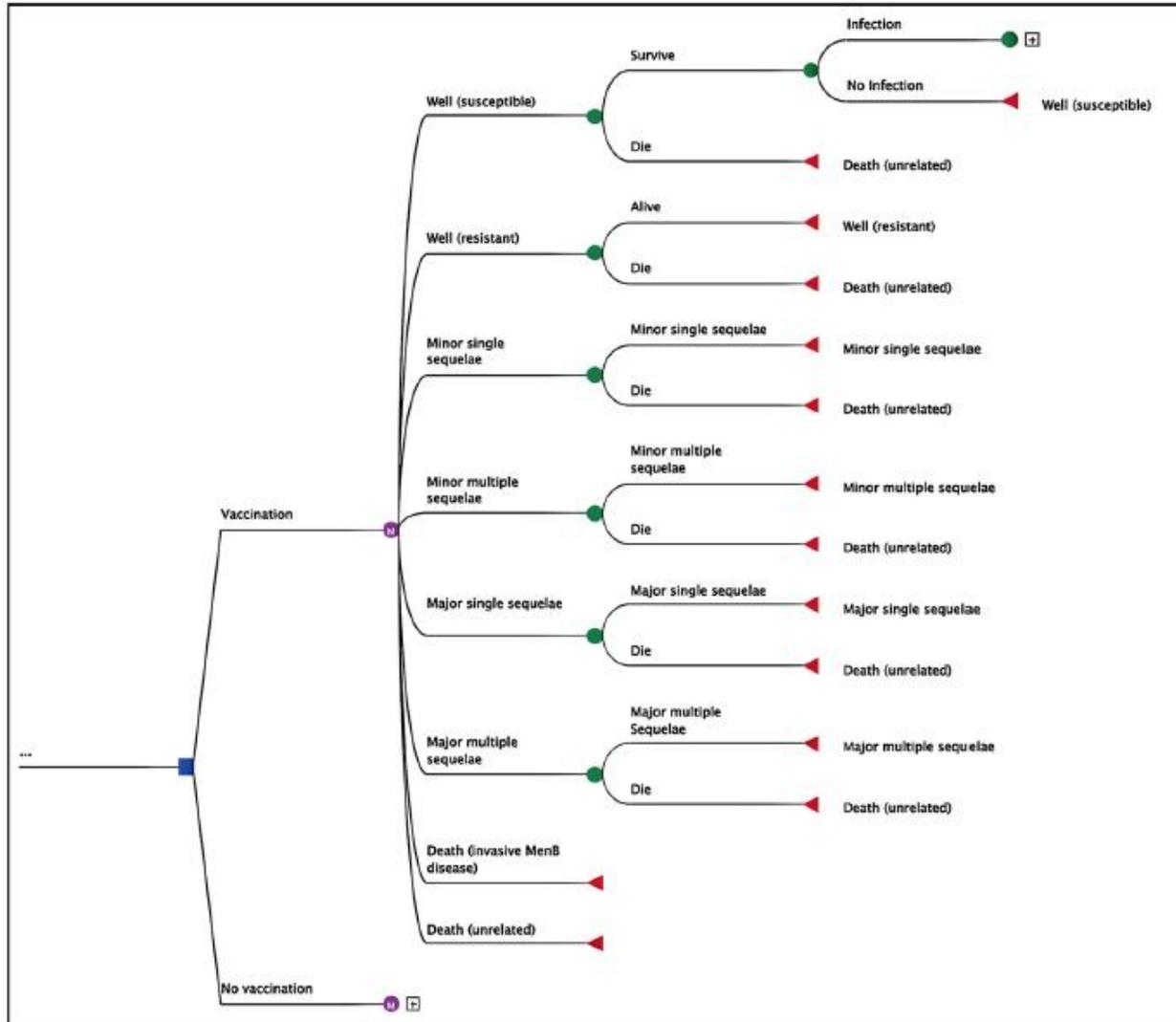
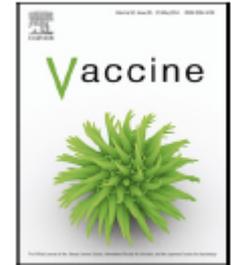
- Transparent in that the structure and implicit or explicit assumptions are all clearly described.
- Static, if vaccination is unlikely to change the force of infection in susceptibles or as a means to make a conservative estimate when externalities from herd-immunity cannot on the whole be adverse.
- Dynamic, if vaccination is likely to change the force of infection in susceptibles, and a static model would not yield a conservative estimate, or if the conservative estimate from a static model does not lead to an outcome which would be considered favourable by decision makers.
- Stochastic if chance plays an important role in the transmission process of the pathogen
- Validated, in as many facets of validation (verification, calibration, face validity, predictive validity) as possible, but at least verified.

Un exemple

Economic evaluation of meningococcal serogroup B childhood vaccination in Ontario, Canada



Hong Anh T. Tu^a, Shelley L. Deeks^{b,c}, Shaun K. Morris^{d,e}, Lisa Striffler^f,
Natasha Crowcroft^{b,c}, Frances B. Jamieson^{b,g}, Jeffrey C. Kwong^{b,c,h,i,j},
Peter C. Coyte^{a,k}, Murray Krahn^{a,j,k,l}, Beate Sander^{a,b,h,k,*}



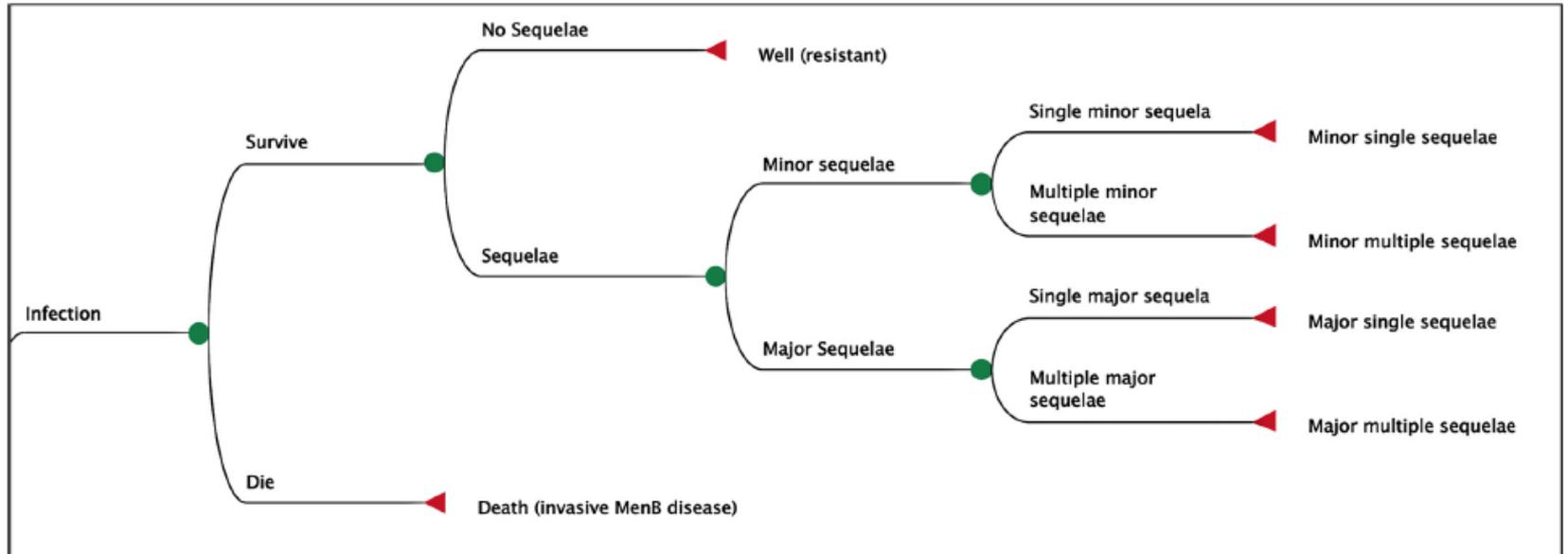


Table 2

Base-case results for MenB vaccination versus no vaccination, undiscounted, discounted at 3% and discounted at 5%.

| | No vaccination | Vaccination | Difference |
|--|----------------|-------------|------------|
| Invasive MenB cases | 23.04 | 18.49 | -4.55 |
| Invasive Men B related deaths | 2.46 | 1.98 | -0.48 |
| QALYs | | | |
| Undiscounted | 11,193,981 | 11,194,020 | 39.02 |
| Discounted at 3% | 4,316,396 | 4,316,411 | 14.93 |
| Discounted at 5% | 2,862,745 | 2,862,755 | 9.76 |
| Cost of vaccination program (CS) | 0 | 46,269,000 | 46,269,000 |
| Cost related to vaccine adverse events (CS) | 0 | 318,383 | 318,383 |
| Treatment costs of invasive MenB cases (CS)^a | | | |
| Undiscounted | 1,491,950 | 1,128,556 | -363,394 |
| Discounted at 3% | 767,055 | 579,935 | -187,119 |
| Discounted at 5% | 631,522 | 480,999 | -150,522 |
| Total cost (CS) | | | |
| Undiscounted | 1,491,950 | 47,715,939 | 46,223,989 |
| Discounted at 3% | 767,055 | 47,167,318 | 46,400,264 |
| Discounted at 5% | 631,522 | 47,068,383 | 46,436,861 |
| ICER (CS/QALY) | | | |
| Undiscounted | | | 1,184,482 |
| Discounted at 3% | | | 3,108,522 |
| Discounted at 5% | | | 4,756,189 |

CS, Canadian Dollar; ICER, Incremental cost-effectiveness ratio; QALY, quality-adjusted life year.

^a Treatment costs include public health cost for contact management as described in Section 2.2.4.

Cost-effectiveness of vaccination against meningococcal B among Dutch infants

Crucial impact of changes in incidence

Koen B. Pouwels,^{1,*} Eelko Hak,¹ Arie van der Ende,^{2,3} Hannah Christensen,⁴ Gemie P.J.M. van den Dobbelsteen^{5,†} and Maarten J. Postma¹

Table 1. Influence of increasing the incidence of meningococcal B disease on the cost-effectiveness of routine infant vaccination with different vaccination strategies

| Vaccination schedule | Base-case incidence* (€/QALY) | Incidence '90-'93* (€/QALY) | Incidence per 100,000 person-years needed to stay below €20,000/QALY ^a | Incidence per 100,000 person-years needed to stay below €50,000/QALY ^a |
|-----------------------|-------------------------------|-----------------------------|---|---|
| 2, 3, 4, 11 mo | 243,778 | 85,931 | 12.58 | 5.72 |
| 2, 3, 4, 11 mo + 12 y | 247,139 | 77,392 | 11.49 | 5.19 |
| 12 + 14 mo | 221,132 | 70,898 | 10.60 | 4.79 |
| 12 + 14 mo + 12 y | 234,548 | 66,811 | 10.08 | 4.53 |

*Base-case incidence was 1.07 per 100,000 person-years; Incidence of '90-'93 was 3.46 per 100,000 person-years. ^aUsing the meningococcal B disease age-distribution of '90-'93.

Vaccinating adolescents against meningococcal disease in Canada: A cost-effectiveness analysis

Philippe De Wals^{a,*}, Laurent Coudeville^b, Pierre Trottier^c, Catherine Chevat^b,
Lonny J. Erickson^d, Van Hung Nguyen^b

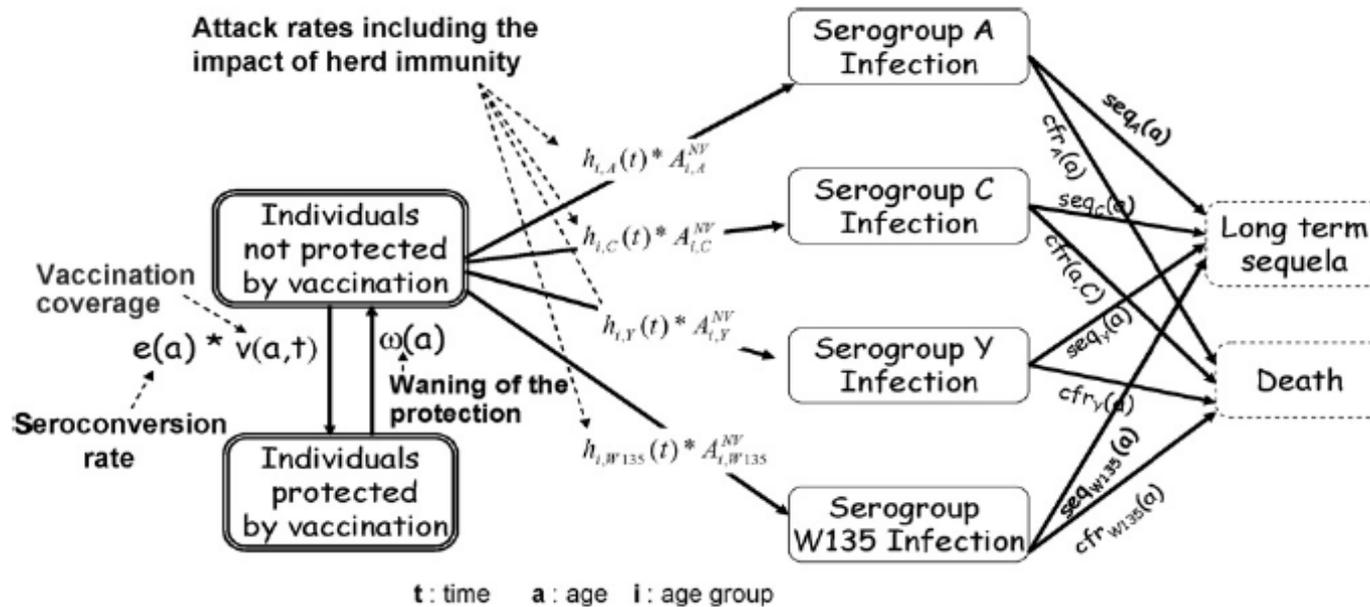


Fig. 1. Schematic representation of the simulation model.

$$h_i(t) = h_0 \sum_j z_{ij} V_j(t) \quad \text{avec} \quad \sum_j z_{ij} = 1 \quad \forall i$$

$$\text{avec} \quad z_{ij} = \frac{a_{ij} c_j N_j}{\sum_k a_{ik} c_k N_k}$$

La paramétrisation du modèle

Herd immunity from meningococcal serogroup C conjugate vaccination in England: database analysis

Mary E Ramsay, Nick J Andrews, Caroline L Trotter, Edward B Kaczmarski, Elizabeth Miller

Attack rate of confirmed meningococcal serogroup C infection in unvaccinated children before and after the launch of the vaccination campaign

| Cohort | July 1998-June 1999 | | | | July 2001-June 2002 | | | | % reduction (95% CI) * | |
|-------------------|---------------------------|-------|------------|----------------------------------|---------------------------|-------|------------------------|----------------------|------------------------|----------------------------------|
| | Date of birth | Cases | Population | Attack rate per 100 000 (95% CI) | Date of birth | Cases | Estimated coverage (%) | Estimated population | | Attack rate per 100 000 (95% CI) |
| Adolescent | 1 Sep 1978 to 31 Aug 1981 | 96 | 1 818 034 | 5.28 (4.2 to 6.3) | 1 Sep 1981 to 31 Aug 1984 | 11 | 66 | 614 110 | 1.79 (0.7 to 2.8) | 66 (37 to 82) |
| School years 7-10 | 1 Sep 1981 to 31 Aug 1985 | 141 | 2 546 938 | 5.54 (4.6 to 6.4) | 1 Sep 1984 to 31 Aug 1988 | 4 | 86 | 359 118 | 1.11 (0.02 to 2.2) | 80 (46 to 93) |
| School years 1-6 | 1 Sep 1985 to 31 Aug 1991 | 76 | 3 911 606 | 1.94 (1.5 to 2.4) | 1 Sep 1988 to 31 Aug 1994 | 5 | 87 | 498 068 | 1.00 (0.1 to 1.9) | 48 (-28 to 79) |
| Preschool | 1 Sep 1991 to 31 Dec 1994 | 81 | 2 055 120 | 3.94 (3.1 to 4.8) | 1 Sep 1994 to 31 Dec 1997 | 6 | 76 | 501 449 | 1.20 (0.2 to 2.2) | 70 (30 to 87) |
| Toddlers | 1 Jan 1995 to 31 Dec 1995 | 41 | 601 045 | 6.82 (4.7 to 8.9) | 1 Jan 1998 to 31 Dec 1998 | 2 | 84 | 97 369 | 2.05 (-0.7 to 4.9) | 70 (-24 to 93) |
| Infants | 1 Jan 1996 to 28 Jul 1996 | 24 | 320 562 | 7.49 (1.5 to 10.5) | 1 Jan 1999 to 28 Jul 1999 | 1 | 80 | 64 112 | 1.56 (-1.5 to 4.6) | 79 (-54 to 97) |
| Overall | | 459 | 11 235 305 | 4.08 (3.7 to 4.5) | | 29 | | 2 134 226 | 1.36 (0.86 to 1.85) | 67 (52 to 77) |

*95% confidence intervals were estimated by using the Taylor series method for relative risks.

range of 48% to 80% across the age groups (table). A smaller fall occurred in adults not eligible for vaccination (aged ≥ 25 years), for whom the incidence declined by 35% (20% to 49%) from 0.53 (193/36 315 726) to 0.34 (123/36 315 726) per 100 000.

Epidemiological impact and cost-effectiveness of vaccination against meningococcal B disease in France

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Base case : impact of vaccination assuming direct effects only (vaccine price = 40 €) – at stationary level

| Vaccination strategy | # cases / year | # cases prevented / y | # deaths prevented / y | Cost per QALY gained (€) |
|--|----------------|-----------------------|------------------------|--------------------------|
| Reference (no vaccination) | 524 | | | - |
| Routine vaccination strategy only | | | | |
| 3, 5, 6 m + 13 m (A) | 432 (-18%) | 92 | 9 | 386,443 |
| 13, 15 m + 27 m (B) | 462 (-12%) | 62 | 6 | 520,110 |
| 15 y (C) | 495 (-5%) | 29 | 2 | 629,488 |
| Routine + Booster + Catch-up strategy | | | | |
| 3,5,6 m+13 m +15y+ C-up (D) | 405 (-23%) | 119 | 11 | 406,889 |
| 13, 15 m+27m +15y+ C-up (E) | 435 (-17%) | 89 | 8 | 522,273 |



Sensitivity analysis: impact of vaccination assuming herd immunity (vaccine price = 40 €) – at stationary level

| Vaccination strategy | # cases / year | # cases prevented / y | # deaths prevented / y | Cost per QALY gained (€) |
|--|----------------|-----------------------|------------------------|--------------------------|
| Reference (no vaccination) | 524 | | | - |
| Routine vaccination strategy only | | | | |
| 3, 5, 6 m + 13 m (A') | 362 (-31%) | 162 | 15 | 225,873 |
| 13, 15 m + 27 m (B') | 391 (-25%) | 133 | 12 | 250,245 |
| 15 y (C') | 397 (-24%) | 127 | 11 | 138,090 |
| Routine + Booster + Catch-up strategy | | | | |
| 3,5,6 m+13 m +15y + C-up (D') | 259 (-51%) | 265 | 25 | 191,366 |
| 13, 15 m+27m +15y + C-up (E') | 283 (-46%) | 241 | 22 | 200,503 |

*Quality adjusted life years

Introducing vaccination against serogroup B meningococcal disease: An economic and mathematical modelling study of potential impact

Hannah Christensen^{a,*}, Matthew Hickman^a, W. John Edmunds^b, Caroline L. Trotter^{a,1}

Re-evaluating cost effectiveness of universal meningitis vaccination (Bexsero) in England: modelling study

OPEN ACCESS

Hannah Christensen *research associate*¹, Caroline L Trotter *senior lecturer*², Matthew Hickman *professor of public health and epidemiology*¹, W John Edmunds *professor of infectious disease modelling*³

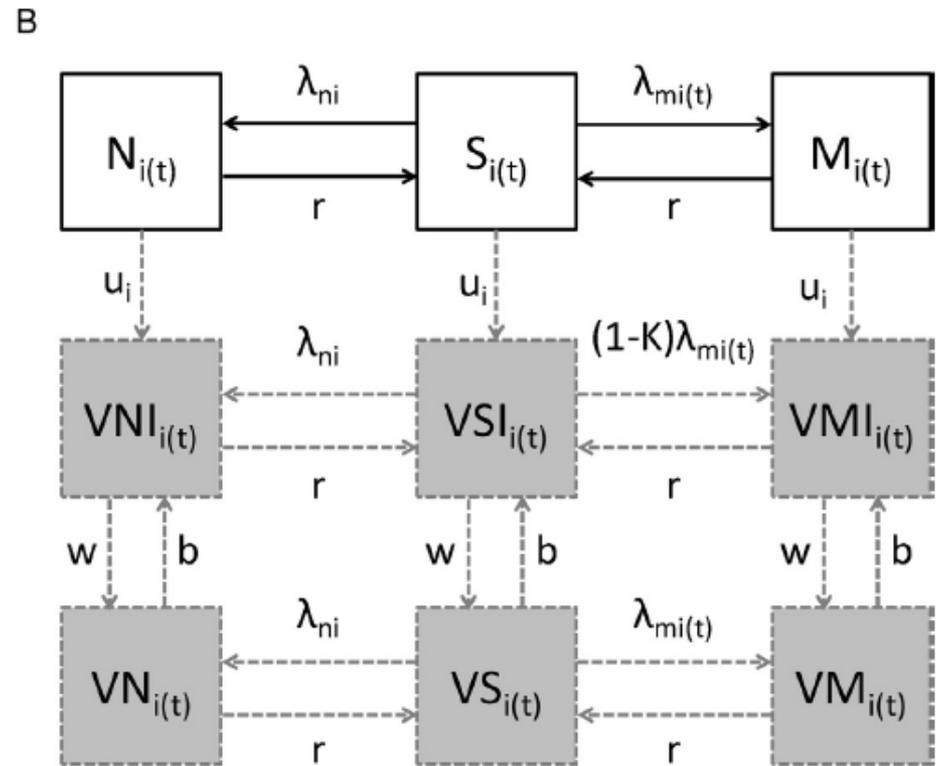
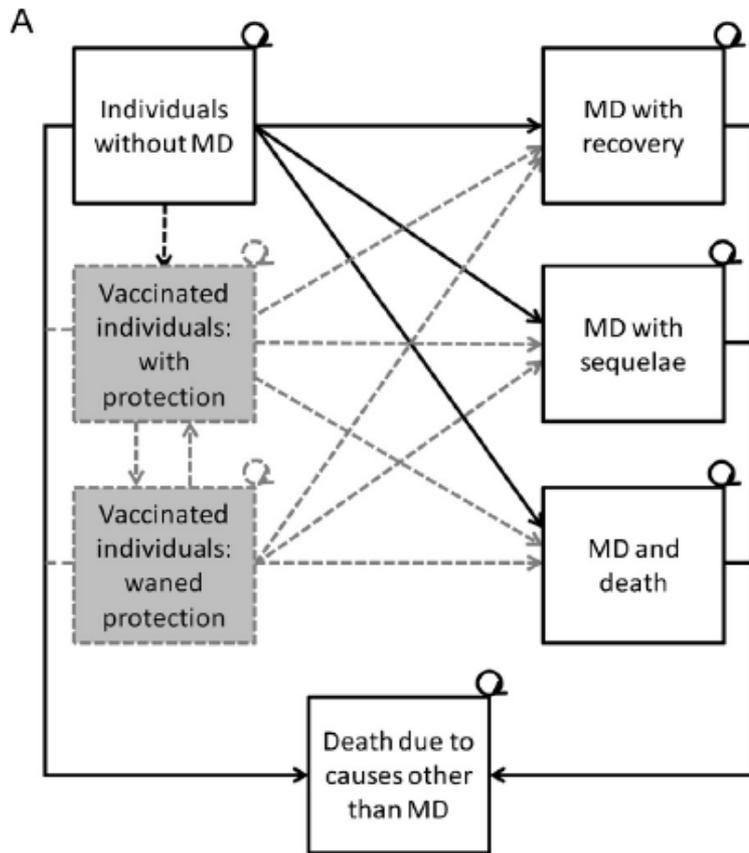


Table 5| Results from dynamic model of cost effectiveness of vaccination (Bexsero) against meningococcal disease. Comparison of vaccination strategies (vaccination v no vaccination) assuming 0% vaccine efficacy against carriage acquisition

| Scenario description | Undiscounted | | | | | | 3.5% discounting for costs and benefits | | 1.5% discounting for costs and benefits | |
|--|---------------|-----------------------------|----------------|------------------|-------------|-------------------------------|---|---|---|---|
| | Cases averted | Cases with sequelae averted | Deaths averted | Life years saved | QALY gained | Net cost of vaccination (£m)* | Cost per QALY gained† | Vaccine price for cost/QALY gained <£20 000 | Cost per QALY gained† | Vaccine price for cost/QALY gained <£20 000 |
| 88% strain coverage | | | | | | | | | | |
| 2, 3, 4, and 12 months | 43 783 | 8 843 | 856 | 41 136 | 140 983 | 19 658.8 | 263 100 | 1 | 181 400 | 5 |
| 2, 3, 4, and 12 months and 13 years | 51 685 | 10 412 | 1 143 | 53 087 | 168 631 | 29 681.9 | 331 600 | NP | 228 500 | 2 |
| 66% strain coverage | | | | | | | | | | |
| 2, 3, 4, and 12 months | 32 837 | 6 632 | 642 | 30 852 | 105 736 | 20 142.8 | 356 100 | NP | 246 800 | 1 |
| 2, 3, 4, and 12 months (with removal of infant meningococcal group C conjugate vaccine cost) | 32 837 | 6 632 | 642 | 30 852 | 105 736 | 19 185.3 | 339 600 | 2 | 235 200 | 5 |
| 2, 4, and 12 months | 32 542 | 6 573 | 636 | 30 573 | 104 774 | 14 757.3 | 265 700 | 1 | 183 300 | 5 |
| 2, 3, 4, and 12 months with 2 dose catch up in 1-4 years | 33 323 | 6 731 | 651 | 31 518 | 108 171 | 20 467.5 | 358 400 | NP | 246 400 | 1 |
| 13 years | 5962 | 1 184 | 217 | 9 008 | 20 850 | 10 100.8 | 927 100 | NP | 627 900 | NP |
| 13 years with 2 dose catch-up in 14-17 years | 6150 | 1 221 | 224 | 9 457 | 21 833 | 10 422.9 | 923 800 | NP | 621 600 | NP |
| 2, 3, 4, and 12 months and 13 years | 38 763 | 7 809 | 857 | 39 815 | 126 473 | 30 245.0 | 447 400 | NP | 309 400 | NP |
| 2, 4, and 12 months and 13 years | 38 468 | 7 749 | 852 | 39 536 | 125511 | 24859.5 | 372 100 | NP | 256 800 | NP |
| 2, 3, 4, and 12 months and 13 years switching after 10 years to 2, 4, and 12 months and 13 years | 38 498 | 7 755 | 852 | 39 583 | 125 670 | 25 397.5 | 394 800 | NP | 266 200 | NP |

Table 4| Results from dynamic model of cost effectiveness of vaccination (Bexsero) against meningococcal disease. Comparison of vaccination strategies (vaccination v no vaccination) assuming 30% vaccine efficacy against carriage acquisition

| Scenario description | Undiscounted | | | | | | 3.5% discounting for costs and benefits | | 1.5% discounting for costs and benefits | |
|--|---------------|-----------------------------|----------------|------------------|-------------|-------------------------------|---|---|---|---|
| | Cases averted | Cases with sequelae averted | Deaths averted | Life years saved | QALY gained | Net cost of vaccination (£m)* | Cost (£)/QALY gained† | Vaccine price (£) for cost/QALY gained <£20 000 | Cost (£)/QALY gained† | Vaccine price (£) for cost/QALY gained <£20 000 |
| 2, 3, 4, and 12 months | 52 152 | 10 513 | 1 117 | 49 503 | 166 812 | 19 309.8 | 221 000 | 3 | 151 400 | 8 |
| 2, 3, 4, and 12 months (with removal of infant meningococcal group C conjugate vaccine cost) | 52 152 | 10 513 | 1 117 | 49 503 | 166 812 | 18 352.3 | 210 500 | 6 | 144 000 | 11 |
| 2, 4 and 12 months | 51 789 | 10 440 | 1 110 | 49 157 | 165 623 | 13 927.3 | 163 100 | 7 | 110 800 | 13 |
| 2, 3, 4, and 12 months with 2 dose catch up in 1-4 years | 53 165 | 10 716 | 1 144 | 50 973 | 171 750 | 19 604.7 | 219 700 | 3 | 149 700 | 8 |
| 13 years | 62 165 | 12 289 | 2 511 | 69 715 | 184 691 | 7946.1 | 104 900 | 14 | 62 100 | 27 |
| 13 years with 2 dose catch up in 14-17 years | 64 667 | 12 783 | 2 613 | 73 998 | 196 544 | 8142.5 | 102 700 | 14 | 60 300 | 28 |
| 2, 3, 4, and 12 months and 13 years | 91 304 | 18 153 | 3 181 | 100 152 | 285 609 | 28 200.3 | 199 000 | 4 | 131 600 | 9 |
| 2, 4 and 12 months and 13 years | 91 118 | 18 116 | 3 178 | 99 953 | 284 931 | 22 810.5 | 163 300 | 6 | 107 300 | 13 |
| 2, 3, 4, and 12 months and 13 years switching after 10 years to 2, 4, and 12 months and 13 years | 91 154 | 18 123 | 3 178 | 100 008 | 285 121 | 23 348.2 | 174 000 | 5 | 111 600 | 13 |

A QUADRIVALENT VACCINE AGAINST SEROGROUP B MENINGOCOCCAL DISEASE: A COST-EFFECTIVENESS STUDY



Federaal Kenniscentrum voor de Gezondheidszorg
Centre Fédéral d'Expertise des Soins de Santé
Belgian Health Care Knowledge Centre

Table 5 – Cost-effectiveness of infant vaccination options, assuming no effect on carriage (compared with no vaccination)

| | Cases averted (%) | Deaths averted (%) | QALYs gained ^a | Net cost of vaccination ^b | ICER: cost per QALY gained ^c |
|--|-------------------|--------------------|---------------------------|--------------------------------------|---|
| <i>Base-case analyses – Infants vaccination at 3, 5, 6 + 12 months</i> | | | | | |
| Routine | 14 (10) | 1 (10) | 65 | €15.6M | €422 700 |
| Partly reimbursed | 13 (9) | 1 (9) | 58 | €22.3M | €663 600 |
| Private market | 5 (4) | 0 (3) | 23 | €8.7M | €667 800 |
| <i>Selected scenario analyses on the routine vaccination strategy</i> | | | | | |
| Increased incidence and case fatality ratios | 34 (10) | 2 | 158 | €14.5M | €167 000 |
| Higher vaccine uptake | 24 (18) | 1 | 110 | 26.9M | €427 400 |
| Higher MATS coverage | 15 (11) | 1 | 70 | €15.5M | €391 100 |
| Alternative assumption for QoL loss for survivors with sequelae | 14 (10) | 1 | 115 | €15.6M | €239 100 |
| All serogroups | 16 (9) | 1 | 77 | €15.5M | €355 000 |
| Best case | 71 (20) | 4 | 284 | €17.0M | €98 300 |

Undiscounted values, except ICER. ICER: incremental cost-effectiveness ratio. M: million. QoL: quality of life. a: QoL lost during the acute phase of the disease and for the adverse events are not included. b: additional cost of vaccination less costs averted through reduction in cases. c: discounted figures rounded to nearest 100.

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Table 6 – Cost-effectiveness of infant, adolescent and combined vaccination options, assuming an effect on carriage (compared with no vaccination)

| | Cases averted (%) | Deaths averted (%) | QALYs gained ^a | Net cost of vaccination ^b | ICER: cost per QALY gained ^c |
|---|-------------------|--------------------|---------------------------|--------------------------------------|---|
| <i>Routine vaccination policy, free of charge</i> | | | | | |
| Infant | 1699 (12) | 85 (12) | 5015 | €1589.5M | €260 700 |
| Adolescent | 8904 (65) | 476 (65) | 21353 | €518.0M | €24 400 |
| Infant + adolescent | 9180 (67) | 490 (67) | 22596 | €2151.7M | €83 000 |
| <i>Partly reimbursed vaccination policy</i> | | | | | |
| Infant | 1532 (11) | 77 (11) | 4525 | €2252.8M | €407 500 |
| Adolescent | 5925 (43) | 317 (43) | 13910 | €496.2M | €34 600 |
| Infant + adolescent | 6676 (49) | 354 (48) | 16360 | €2772.3M | €146 300 |
| <i>Private market vaccination policy</i> | | | | | |
| Infant | 604 (4) | 30 (4) | 1788 | €877.3M | €402 000 |
| Adolescent | 2090 (15) | 112 (15) | 4992 | €198.1M | €37 700 |
| Infant + adolescent | 2592 (19) | 137 (19) | 6499 | €1076.4M | €142 800 |

Undiscounted values, except ICER. ICER: incremental cost-effectiveness ratio. M: million. a: QoL lost during the acute phase of the disease and for the adverse events are not included. b: additional cost of vaccination less costs averted through reduction in cases. c: discounted figures rounded to nearest 100.

Impact of MenBvac, an outer membrane vesicle (OMV) vaccine, on the meningococcal carriage



Valérie Delbos^{a,b,*}, Ludovic Lemée^{a,b}, Jacques Bénichou^{a,c}, Gilles Berthelot^d,
Ala-Eddine Deghmane^e, Jean-Philippe Leroy^a, Estelle Houivet^a, Eva Hong^e,
Muhammed-Kheir Taha^{e,1}, François Caron^{a,b,1}, on behalf of the B14 STOP study group²

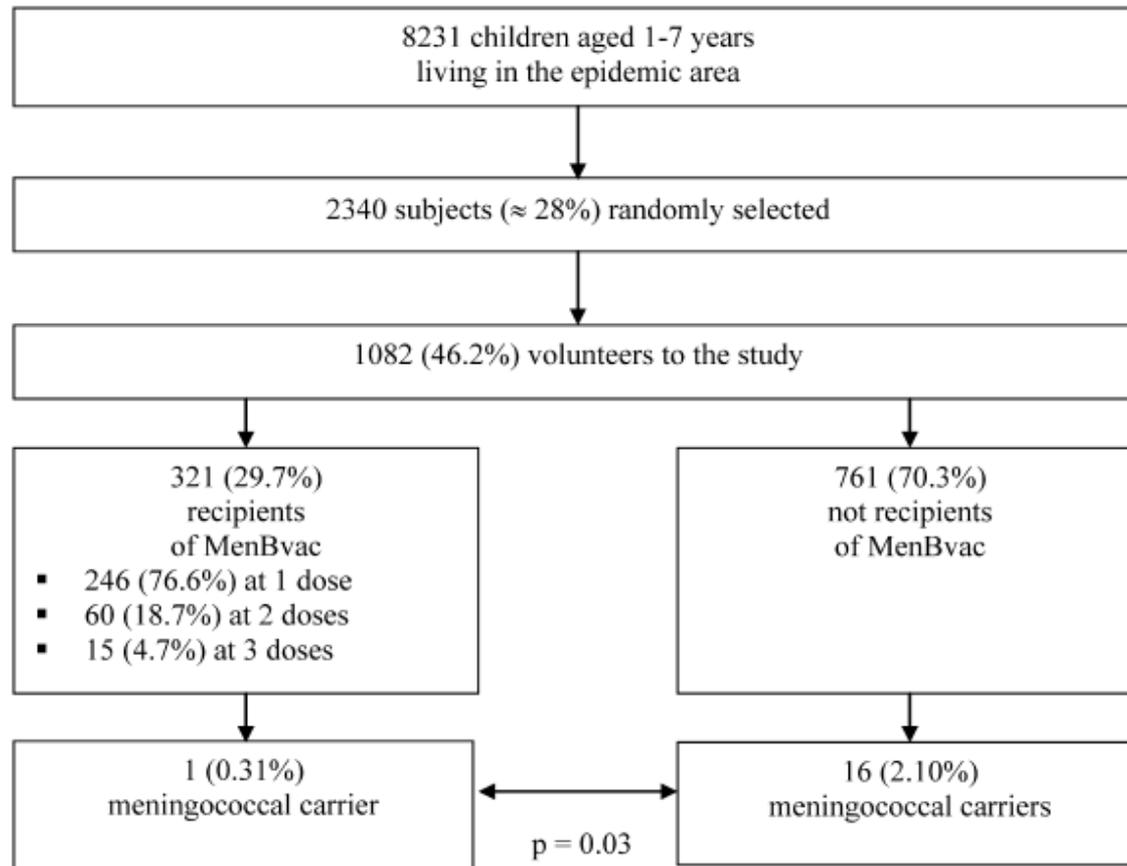


Fig. 1. Flow-chart of the study and main results.

Les difficultés

- Structure mathématique du modèle (→ propriétés: convergence, équilibre)
- Calibration, validation du modèle
- Utilisation recommandée de logiciels spécifiques (mais utilisation de feuilles de calcul possible)
- Difficulté à conduire les analyses de sensibilité probabilistes

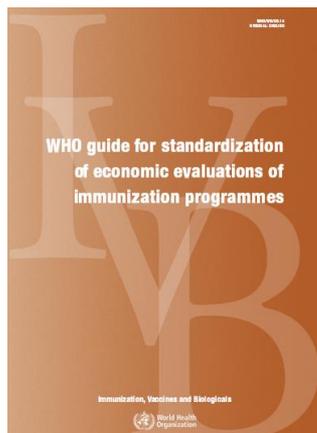


Table 8: Practical differences of static versus dynamic models for economic evaluation

| | Static | Dynamic |
|---|---|--|
| Typical population in which costs and effects are monitored | A single ageing cohort (with removal of deaths from all causes through time) | The entire population (with introductions of births and removal of deaths from all causes through time) |
| Development complexity | Easy to develop, embedded in traditional health economic methods | Not part of the traditional toolbox of epidemiologists and health economists |
| Ready to use software | Spreadsheets, e.g. MS Excel™ (www.microsoft.com) @Risk™ (www.palisade.com) TreeAge Pro™ (www.treeage.com), Crystal Ball™ (www.crystalball.com) | Berkeley Madonna™ (www.berkeleymadonna.com) Model Maker™ (www.modelkinetix.com) Stella™ (www.iseesystems.com) Vensim™ (www.vensim.com) |
| Required data | Requires (usually age-specific) data on epidemiology, demography, course of illness, vaccine efficacy, costs | Same as with static models + average duration of infectiousness + information on relevant mixing patterns between infectious and susceptible people |

La structure d'un modèle dynamique

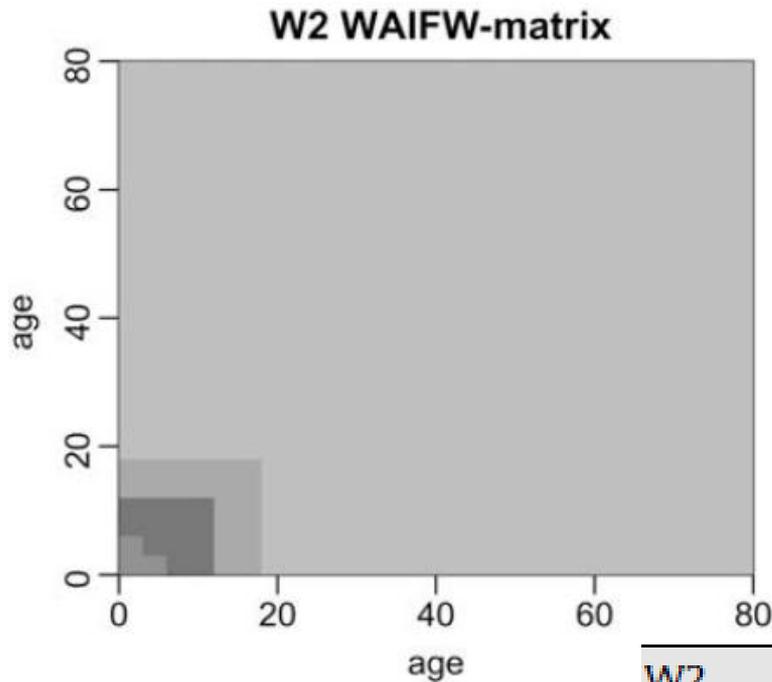
$$\frac{\partial S(a, t)}{\partial a} + \frac{\partial S(a, t)}{\partial t} = -S(a, t) \cdot \underbrace{\int_0^{\infty} \beta(a, a') \cdot I(a', t) da'}_{\lambda(a, t)} - \mu(a, t) \cdot S(a, t)$$

$$\begin{aligned} \frac{\partial L(a, t)}{\partial a} + \frac{\partial L(a, t)}{\partial t} &= +S(a, t) \cdot \underbrace{\int_0^{\infty} \beta(a, a') \cdot I(a', t) da'}_{\lambda(a, t)} - \mu(a, t) \cdot L(a, t) \\ &\quad - e(a, t) \cdot L(a, t) \end{aligned}$$

$$\frac{\partial I(a, t)}{\partial a} + \frac{\partial I(a, t)}{\partial t} = e(a, t) \cdot L(a, t) - \mu(a, t) \cdot I(a, t) - g(a, t) \cdot I(a, t)$$

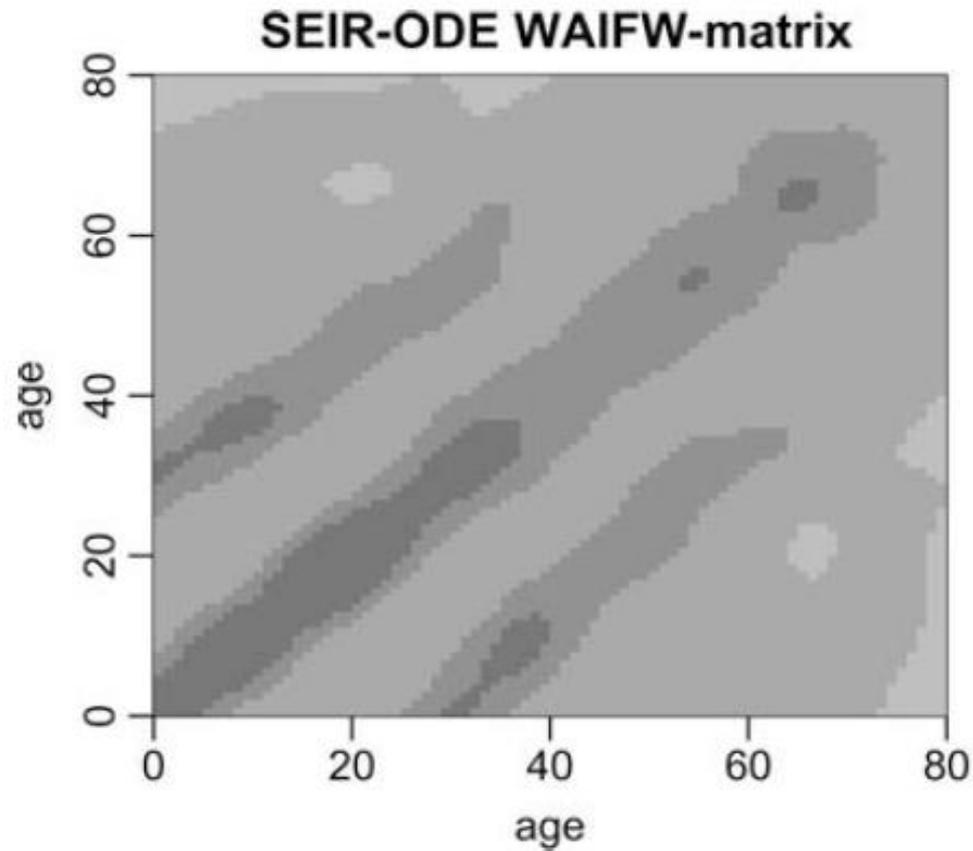
$$\frac{\partial R(a, t)}{\partial a} + \frac{\partial R(a, t)}{\partial t} = g(a, t) \cdot I(a, t) - \mu(a, t) \cdot R(a, t)$$

La matrice de contact (1)



| W2 | [0.5 3[| [3 6[| [6 12[| [12 18[| [18 26[| [26 80[|
|---------|-----------|-----------|-----------|-----------|-----------|-----------|
| [0.5-3[| β_1 | β_1 | β_3 | β_4 | β_5 | β_6 |
| [3 6[| β_1 | β_2 | β_3 | β_4 | β_5 | β_6 |
| [6 12[| β_3 | β_3 | β_3 | β_4 | β_5 | β_6 |
| [12 18[| β_4 | β_4 | β_4 | β_4 | β_5 | β_6 |
| [18 26[| β_5 | β_5 | β_5 | β_5 | β_5 | β_6 |
| [26 80[| β_6 | β_6 | β_6 | β_6 | β_6 | β_6 |

La matrice de contact (2)



Etude COMES-F (1)

- **Rationnel/Objectif :**

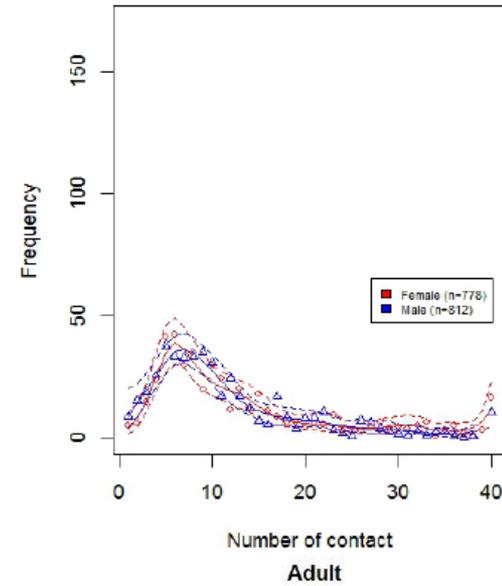
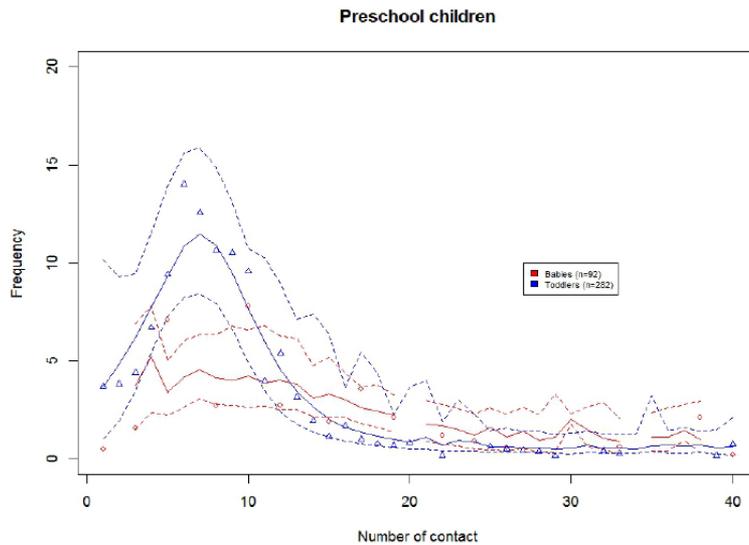
- Utilisation de plus en plus fréquente de modèles mathématiques pour l'évaluation des mesures de **prévention et de contrôle des maladies transmissibles**
- Nécessité de connaître la **structure des contacts au sein de la population** (entre classes d'âge, entre classes de risque...) pour estimer la force de l'infection
- Deux approches pour estimer la matrice WAIFW (« Who Acquires Infection from Whom ») :
- La structure de la matrice est fixée a priori, les paramètres sont estimés lors de la calibration du modèle (/ données d'incidence, de séroprévalence...). Différentes structures de la matrice peuvent être successivement testées [Anderson et May (1991)]
- La structure de la matrice est fournie par l'observation des contacts au sein de la population. La calibration du modèle permet d'estimer la(les) probabilité(s) de transmission [paramètre(s) q]
- Etudes conduites dans de nombreux pays européens [Mossong et al. (2008)]
- **Pas de données disponibles pour la France**

Etude COMES-F (2)

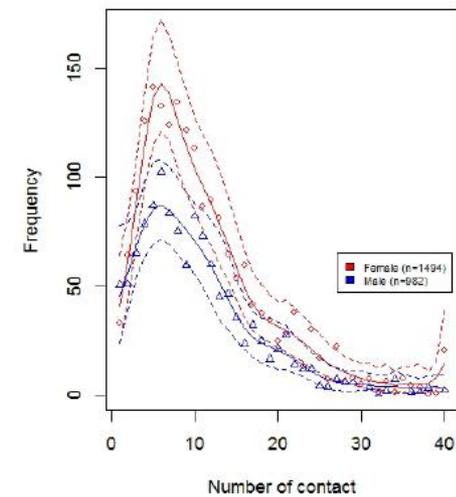
- **Résultats :**
 - Nombre moyen de contacts :
 - 9,5 / personne / jour en excluant les contacts des personnes ayant de nombreux contacts professionnels
 - 13,9 / personne / jour en prenant en compte de tous les contacts
 - Des contacts majoritairement « assortifs » avec un effet de la parentalité

Etude COMES-F (3)

Children

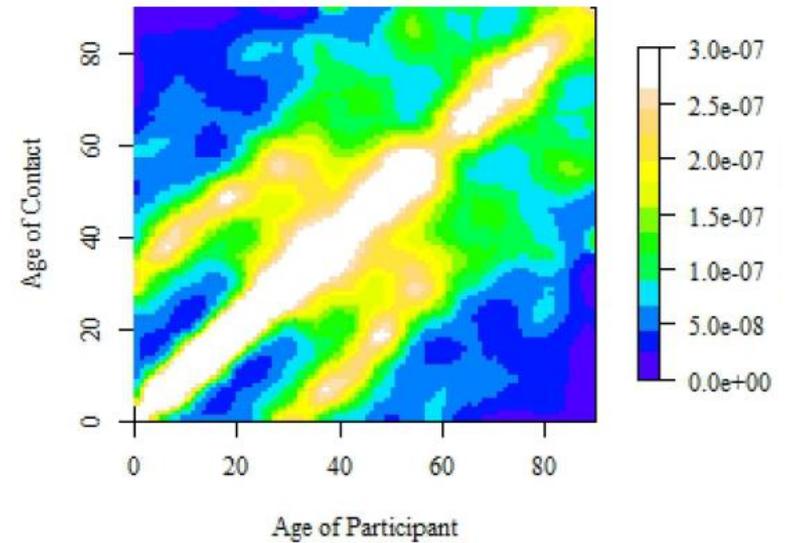
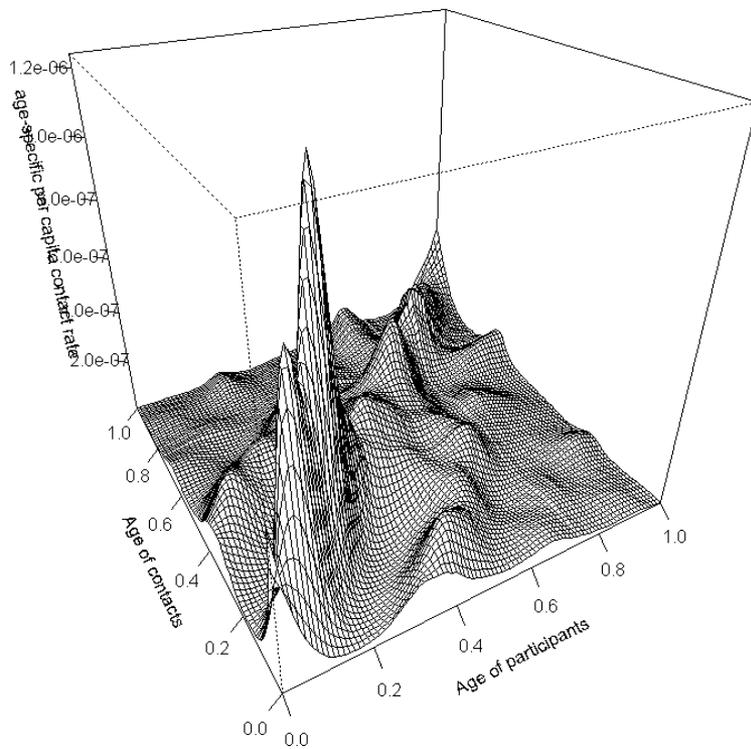


Adult



Béraud et al. (2015)

Etude COMES-F (4)



Béraud et al. (2015)

Etude COMES-F (5)

- **Résultats (suite) :**
 - Variables influençant le nombre de contacts (NRC = nombre relatif de contacts) :
 - La vague d'enquête
 - L'âge
 - Le sexe (NRC hommes : 0,92 (0,86-0,99))
 - Le moment
 - NRC week-end : 0,79 (0,73-0,86)
 - NRC vacances scolaires : 0,79 (0,74-0,84)

Merci de votre attention

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