

Cette présentation a été effectuée le 2 décembre 2024, au cours de la journée « Contrôle des BGNPC au Québec : c'est le temps d'agir! » dans le cadre des 27es Journées annuelles de santé publique.

Épidémiologie et impacts cliniques des BGNPC

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AMMI Canada



OBJECTIFS

1. Décrire l'état de la situation et l'évolution des **BGNPC** dans les milieux de soins du Québec.
2. Évaluer le **fardeau** de la résistance aux antimicrobiens (**RAM**) et des **EPC** au **Canada** et dans le **monde** à l'aide d'une gamme d'indicateurs
3. Comprendre **l'impact clinique** des infections à EPC

Déclaration de conflit d'intérêts

Yves Longtin

Relations avec les organismes sans but lucratif :

Salary support:

Fonds de recherche en santé du Québec

Scientific committee membership:

WHO Working Group on Antimicrobial Resistance

AMMI Canada

Comité des infections nosocomiales du Québec

Canadian Nosocomial Infection Surveillance Program

Qu'est-ce que la RAM ?

- Conceptuellement:
 - La résistance aux antimicrobiens (RAM) se produit lorsque les bactéries, les virus, les champignons et les parasites **ne réagissent plus aux médicaments antimicrobiens**
- Conséquences:
 - Les antimicrobiens deviennent **inefficaces**
 - Les infections deviennent difficiles ou **impossibles à traiter**
- La RAM est un **processus naturel** ! mais **s'accélère** par l'activité humaine telle que la surutilisation de l'ATB

<https://www.who.int/news-room/fact-sheets/detail/antimicrobial-resistance>

L'histoire des antibiotiques n'a-t-elle commencé qu'en 1928 ?



Sir Alexander Fleming (1881 – 1955)

Chronologie des antibiotiques (et de la RAM)

Daptomycin
30 million years

Vancomycin
240 million years

Streptomycin
610 million years

Erythromycin
880 million years

Beta-lactamases
> 2 billion years

Original carbapenem
thienamycin

Source: *Streptomyces cattleya*

Beta-lactams *Aspergillus*,
Streptomyces, *Flavobacterium*
and *Burkholderia*



Wright (2007). *Nat Rev Microbiol* 5:175-186

Hall et Barlow (2004). *Drug Resist Updat* 7:111-23

Organismes multirésistants

S. aureus résistant à la méthicilline (SARM)

Entérobactéries résistantes aux carbapénèmes

Acinetobacter antibiorésistant

Entérocoque résistant à la vancomycine (E

NEW!

Candida auris

“Has-beens”

Entérocoque résistant à la gentamicine

Entérobactéries productrices de B-lactamase à spectre étendu (ESBL)



Global burden of bacterial antimicrobial resistance in 2019: a systematic analysis



Antimicrobial Resistance Collaborators*



Summary
Background Antimicrobial resistance (AMR) poses a major threat to human health around the world. Previous publications have estimated the effect of AMR on incidence, deaths, hospital length of stay, and health-care costs for specific pathogen–drug combinations in select locations. To our knowledge, this study presents the most comprehensive estimates of AMR burden to date.

Lancet 2022; 399: 629–55
Published Online
January 20, 2022
[https://doi.org/10.1016/S0140-6736\(21\)02724-0](https://doi.org/10.1016/S0140-6736(21)02724-0)

- Objectif : Quantifier le fardeau de la RAM dans 204 pays en 2019
 - 12 syndromes infectieux et 23 agents pathogènes
 - Sources multiples de données
 - Rev syst, bases de données de surveillance, bases de données hospitalières
 - Total: 471 million de données
- 2 comparaisons
 - Comparaison de l’infection due à la RAM et à la même infection, mais sans RAM
 - Comparaison de l’infection due à la RAM à l’absence d’infection
 - Cela permet de comparer le fardeau de l’infection et le fardeau de la RAM



Global burden of bacterial antimicrobial resistance in 2019: a systematic analysis

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- **Conclusions**
 - Décès associés à RAM: 4.95 million (95% CI, 3.62–6.57)
- **Décès attribuables à RAM: 1.27 million (95% CI, 0.91–1.71)***
- Gains découlant de la prévention des infections et de la prévention de la propagation de la RAM
- Seulement 1 RAM majeure est évitable par la vaccination (pneumocoque)

*Comparateurs : Décès dus au VIH : 680K Malaria : 627k

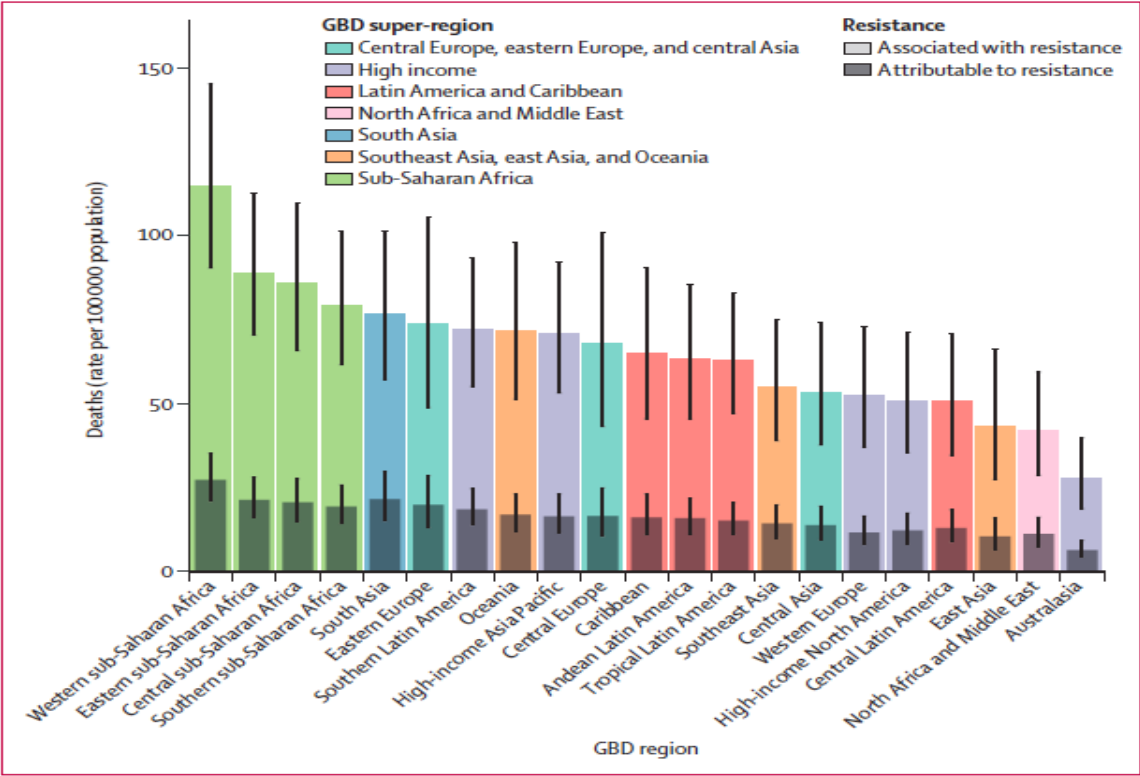


Figure 2: All-age rate of deaths attributable to and associated with bacterial antimicrobial resistance by GBD region, 2019
 Estimates were aggregated across drugs, accounting for the co-occurrence of resistance to multiple drugs. Error bars show 95% uncertainty intervals. GBD=Global Burden of Diseases, Injuries, and Risk Factors Study.

Global burden of bacterial antimicrobial resistance in 2019: a systematic analysis

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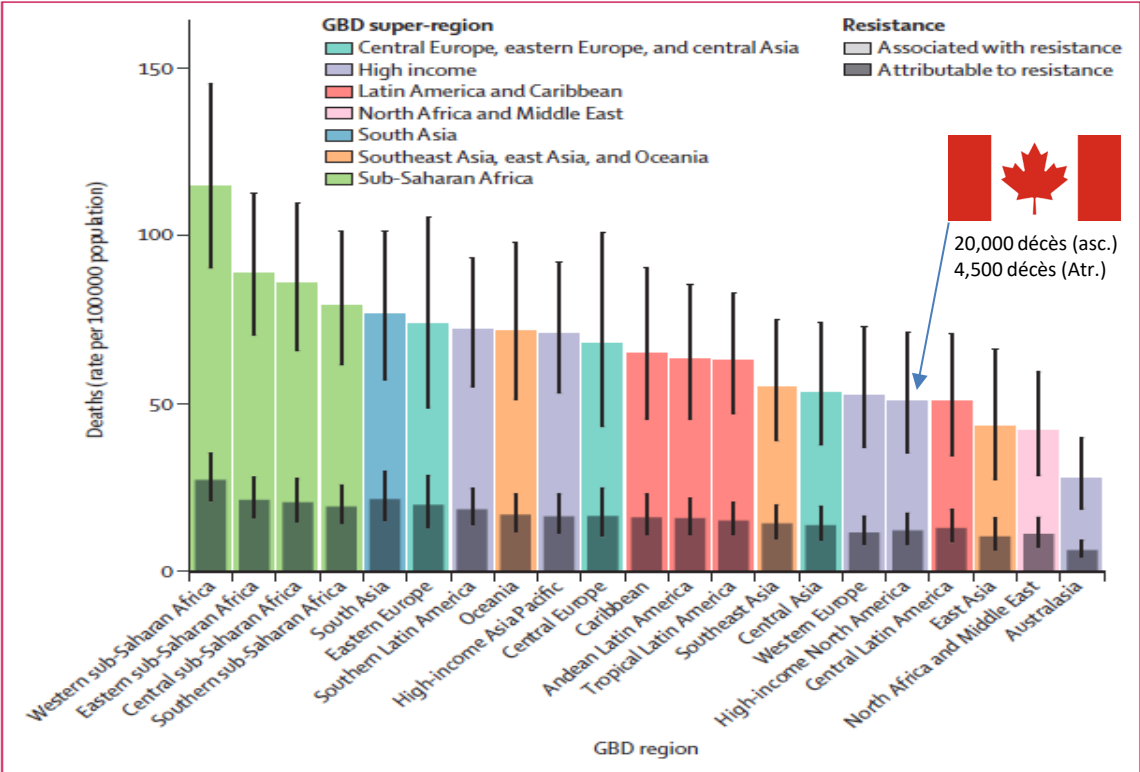


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Causes de décès au Canada

Top 12 leading causes of death in Canada

Cause	No. deaths (2019)
Malignant neoplasms	80372
Diseases of heart	53364
Accidents (unintentional injuries)	15527
Cerebrovascular diseases	13717
Chronic lower respiratory diseases	12902
Diabetes mellitus	6987
Influenza and pneumonia	6945
Alzheimer's disease	6181
Intentional self-harm (suicide)	4581
Chronic liver disease and cirrhosis	3708
Nephritis, nephrotic syndrome and nephrosis	3770
Other ill-defined and unspecified causes of mortality	3378
Total, all causes of death	285301

20 000 décès
associés à RAM



4,500 décès
Attributable
À RAM



Prévisions, 2050

	Associated death counts (thousands)				Associated death rates per 100 000				Attributable death counts (thousands)				Attributable death rate per 100 000			
	1990	2019	2021	2050	1990	2019	2021	2050	1990	2019	2021	2050	1990	2019	2021	2050
Global	4780 (4000–5550)	4940 (4430–5450)	4710 (4230–5190)	8220 (6850–9650)	89.6 (75.0–104)	63.8 (57.2–70.4)	59.7 (53.6–65.7)	87.7 (73.2–104)	1060 (841–1270)	1200 (1050–1350)	1140 (1000–1280)	1910 (1560–2260)	19.8 (15.8–23.9)	15.5 (13.6–17.4)	14.5 (12.7–16.2)	20.4 (16.6–24.2)
Central Europe, Eastern Europe, and Central Asia	285 (240–330)	281 (252–310)	265 (235–295)	360 (297–420)	67.8 (57.1–78.4)	67.1 (60.1–74.0)	63.4 (56.2–70.6)	90.5 (76.3–106)	63.0 (49.5–76.6)	68.6 (59.2–78.0)	64.0 (55.2–72.8)	82.9 (67.1–98.7)	15.0 (11.8–18.2)	16.4 (14.1–18.6)	15.3 (13.2–17.4)	20.8 (17.0–24.8)
High-income	477 (385–568)	579 (513–644)	553 (489–618)	883 (674–1040)	52.5 (42.4–62.5)	53.3 (47.3–59.3)	50.7 (44.8–56.6)	78.1 (59.8–92.2)	108 (82.9–133)	131 (115–146)	125 (110–140)	192 (146–225)	11.9 (9.12–14.7)	12.0 (10.6–13.5)	11.4 (10.1–12.8)	17.0 (12.9–19.9)
Latin America and Caribbean	247 (210–284)	339 (305–372)	322 (285–360)	650 (520–808)	63.3 (53.8–72.8)	57.9 (52.2–63.6)	54.2 (47.9–60.6)	96.7 (78.0–119)	55.7 (44.4–67.0)	82.4 (72.4–92.5)	78.1 (67.7–88.6)	148 (117–185)	14.3 (11.4–17.2)	14.1 (12.4–15.8)	13.2 (11.4–14.9)	22.1 (17.5–27.2)
North Africa and Middle East	264 (219–310)	243 (211–274)	226 (194–259)	525 (430–641)	78.0 (64.5–91.4)	40.0 (34.8–45.2)	36.3 (31.1–41.5)	61.4 (49.2–73.9)	63.0 (49.4–76.7)	64.7 (54.4–75.1)	60.2 (50.1–70.3)	133 (107–161)	18.6 (14.6–22.6)	10.7 (8.97–12.4)	9.66 (8.05–11.3)	15.5 (12.3–19.0)
South Asia	1400 (1170–1630)	1350 (1200–1500)	1260 (1110–1420)	2400 (1910–2980)	128 (107–149)	74.7 (66.5–83.0)	68.5 (60.2–76.8)	114 (91.3–141)	308 (249–367)	356 (303–408)	335 (282–387)	604 (463–743)	28.2 (22.8–33.6)	19.7 (16.8–22.6)	18.1 (15.3–21.0)	28.8 (22.5–35.6)
Southeast Asia, East Asia, and Oceania	1110 (916–1310)	1140 (995–1290)	1150 (1010–1290)	1940 (1580–2370)	65.8 (54.2–77.5)	52.8 (46.0–59.5)	52.7 (46.3–59.1)	92.0 (75.0–113)	250 (195–304)	271 (235–307)	270 (237–304)	428 (348–511)	14.8 (11.5–18.0)	12.5 (10.9–14.2)	12.4 (10.8–13.9)	20.3 (16.4–24.5)
Sub-Saharan Africa	990 (797–1180)	1010 (811–1200)	923 (732–1110)	1470 (1140–1860)	202 (162–241)	93.2 (75.1–111)	81.5 (64.6–98.4)	69.3 (53.7–87.5)	210 (158–262)	227 (179–276)	209 (161–257)	323 (245–416)	42.8 (32.2–53.3)	21.0 (16.5–25.5)	18.5 (14.2–22.7)	15.3 (11.5–19.5)

Table 3: Deaths (in counts and all-age rates) associated with and attributable to bacterial antimicrobial resistance, globally, by GBD super-region, for 1990, 2019, 2021 and 2050



Prévisions 2050

Décès associés à la
RAM : 37 000

RAM Décès
attribuables : 8 000

Prévisions démographiques : 48M

Global burden of bacterial antimicrobial resistance 1990–2021: a systematic analysis with forecasts to 2050

GBD 2021 Antimicrobial Resistance Collaborators*

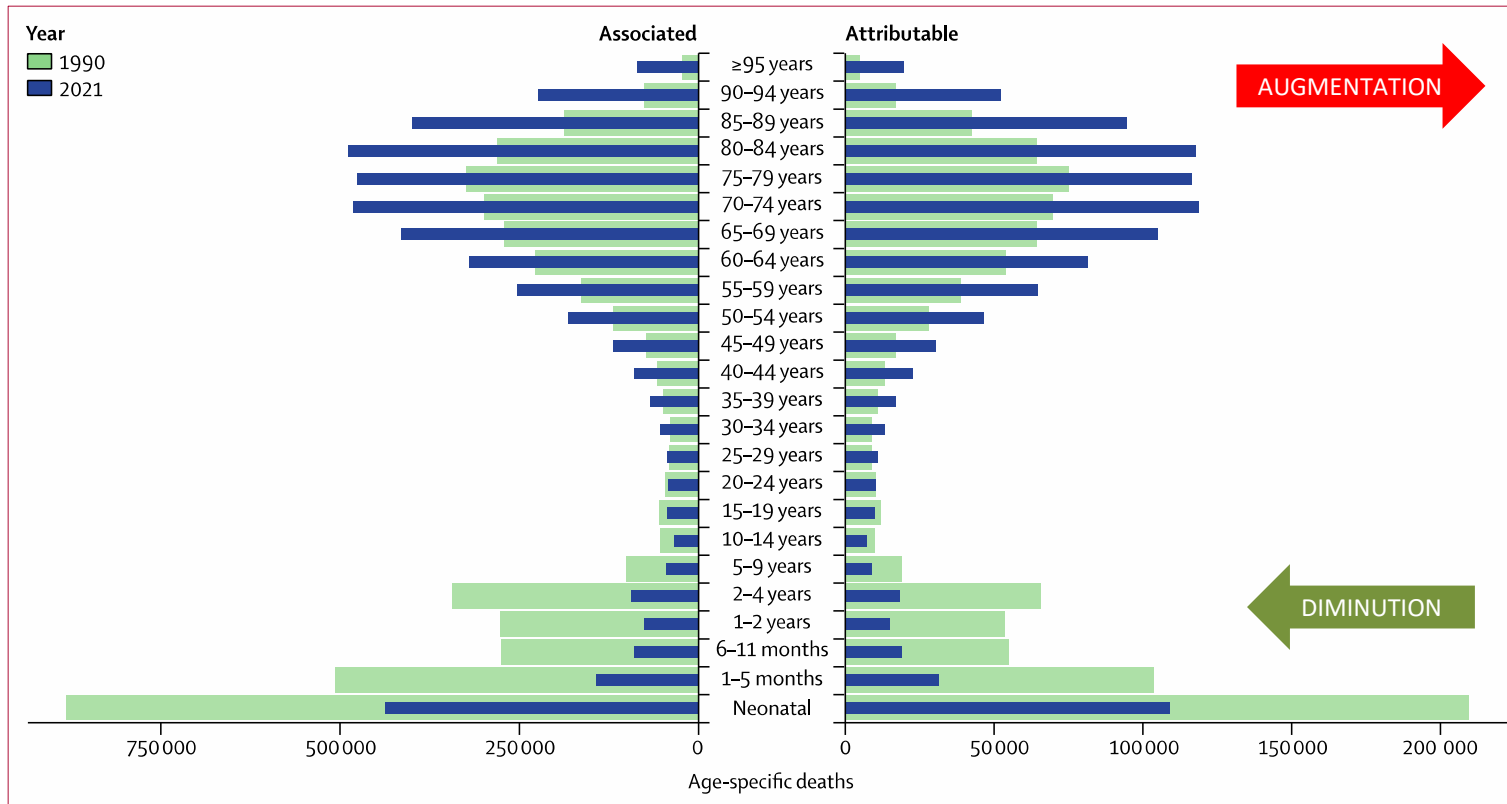


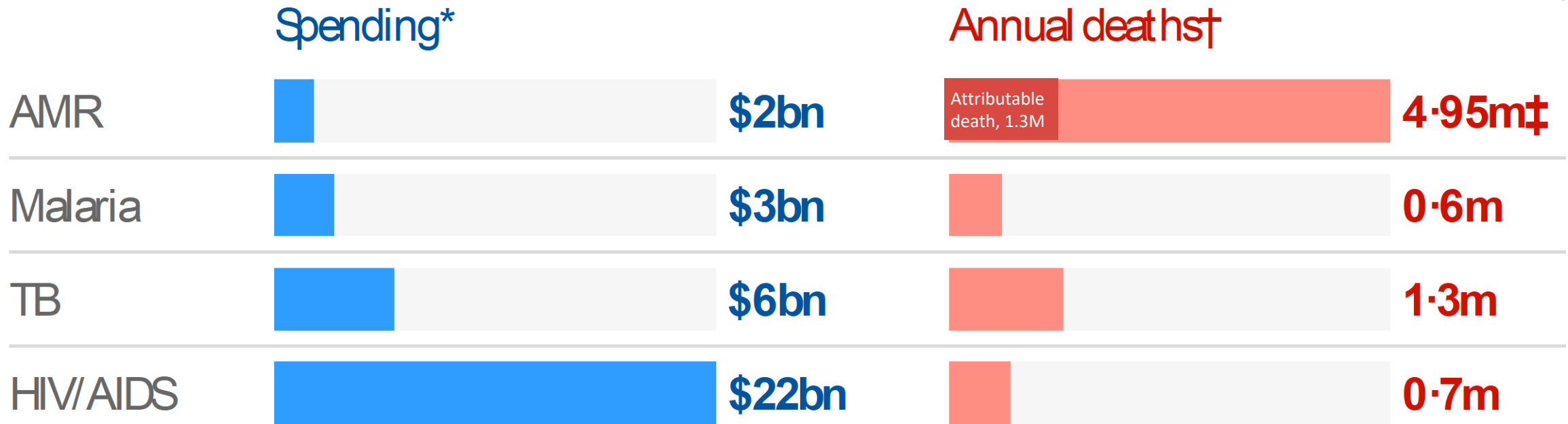
Figure 2: Deaths attributable and associated with antimicrobial resistance, by detailed age group, for 1990 and 2021
Counterfactuals have distinct x-axes.

Diminution
des décès liés
à la RAM
chez les
enfants, mais
augmentation
du nombre
de personnes
> 50



Antibiotics

The ability to deploy drugs and diagnostics for HIV/AIDS, malaria, and TB is well developed. However, similar support is absent for antibiotics even though non-TB bacterial infections kill far more than HIV/AIDS, malaria, and TB combined.



*Average spending, 2017 to 2021; †Estimated global deaths in 2019; ‡Associated deaths

Political Declaration of the High-level Meeting on Antimicrobial Resistance

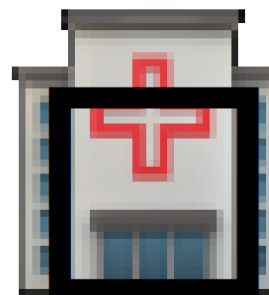
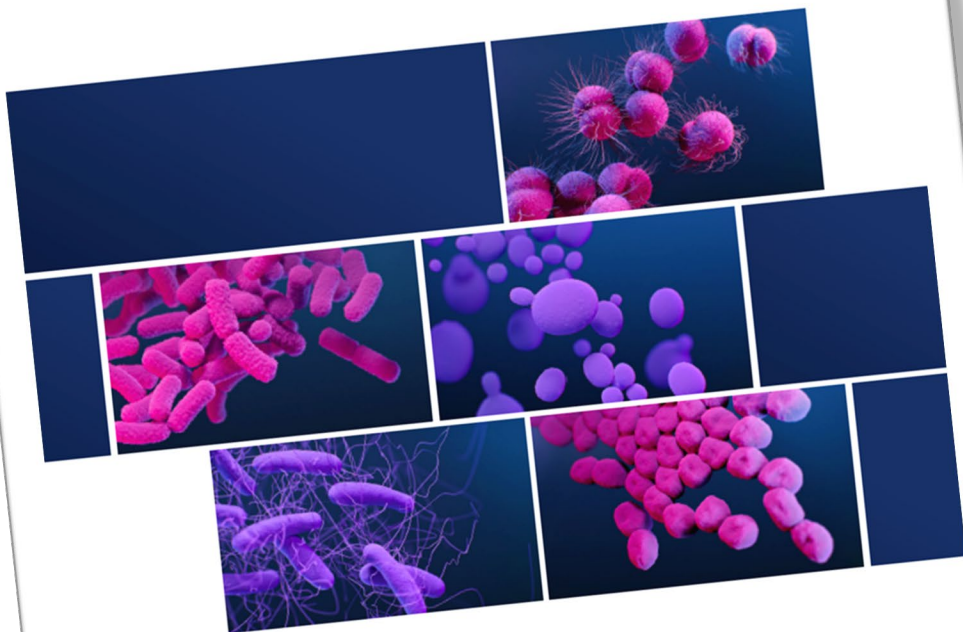
Sept 9th 2024

- AMR is one of the most urgent global health threats
- AMR demands immediate action to safeguard our ability to treat human, animal and plant disease and enhance food safety
- Globally, AMR could result in USD 1 trillion of additional health-care costs by 2050
- AMR can lead to 11% decline in livestock production by 2050
- Commit to reduce global deaths associated with AMR by 10% by 2030 compared to their 2019 levels.

La RAM est-elle un problème lié aux communautés ou aux milieux de soins de santé ?



ANTIBIOTIC RESISTANCE THREATS IN THE UNITED STATES 2019



48% (10/21)
**Healthcare
Associated**

52%
**Community
Associated**

 Mainly a threat in healthcare settings

Urgent Threats

-  Carbapenem-resistant *Acinetobacter*
-  *Candida auris*
-  *Clostridioides difficile*
-  Carbapenem-resistant Enterobacteriaceae
-  Drug-resistant *Neisseria gonorrhoeae*

CPE

Serious Threats

- Drug-resistant *Campylobacter*
-  Drug-resistant *Candida*
-  ESBL-producing Enterobacteriaceae
-  Vancomycin-resistant *Enterococci*
-  Multidrug-resistant *Pseudomonas aeruginosa*
- Drug-resistant nontyphoidal *Salmonella*
- Drug-resistant *Salmonella* serotype Typhi
- Drug-resistant *Shigella*
-  Methicillin-resistant *Staphylococcus aureus*
- Drug-resistant *Streptococcus pneumoniae*
- Drug-resistant Tuberculosis

VRE

MRSA

Concerning Threats

- Erythromycin-resistant group A *Streptococcus*
- Clindamycin-resistant group B *Streptococcus*

Watch List

-  Azole-resistant *Aspergillus fumigatus*
- Drug-resistant *Mycoplasma genitalium*
- Drug-resistant *Bordetella pertussis*

Les 15 principaux agents pathogènes nosocomiaux NHSN, adultes, 2018 - 2021

Pathogen	# Pathogens	% Pathogens	Rank	Emerging AMR
<i>Escherichia coli</i>	73,556	16.2	1	Yes (AmpC, ESBL, CPE)
<i>Staphylococcus aureus</i>	51,131	11.3	2	Yes (MRSA)
<i>Enterococcus faecalis</i> ²	39,129	8.6	3	Partly (VRE)
Select <i>Klebsiella</i> spp.	38,496	8.5	4	Yes (CPE)
<i>Pseudomonas aeruginosa</i>	36,004	7.9	5	Yes (CRO)
CoN staphylococci	32,276	7.1	6	
<i>Enterobacter</i> spp.	18,431	4.1	7	Yes (CPE)
<i>Enterococcus faecium</i> ²	16,904	3.7	8	Yes (VRE)
<i>Candida albicans</i> ²	16,458	3.6	9	
<i>Proteus</i> spp.	13,953	3.1	10	Yes (CPE)
<i>Bacteroides</i> spp.	11,602	2.6	11	
Viridans group streptococci	9,962	2.2	12	
Other <i>Candida</i> spp. ²	9,803	2.2	13	Yes (<i>C. auris</i>)
Other <i>Enterococcus</i> spp. ²	9,091	2.0	14	
<i>Candida glabrata</i> ²	7,622	1.7	15	
Other pathogens	68,522	15.1		
Total	452,940	100.0		

Global R to 3GC: 42%
Global MRSA: 35%

WHO Glass 2022



La prévention
de la
transmission
et des
infections
nosocomiales
est essentiel

Portrait mondial de la RAM

Global Antimicrobial
Resistance and Use
Surveillance System
(GLASS) Report 2022

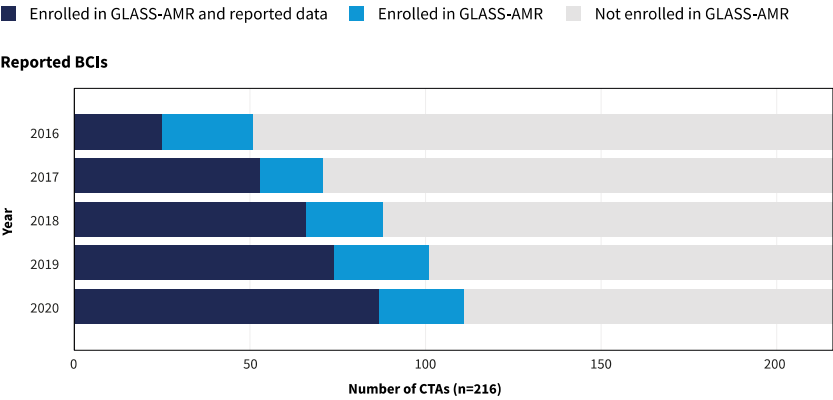


<https://iris.who.int/bitstream/handle/10665/364996/9789240062702-eng.pdf?sequence=1>



GLASS, WHO 2022

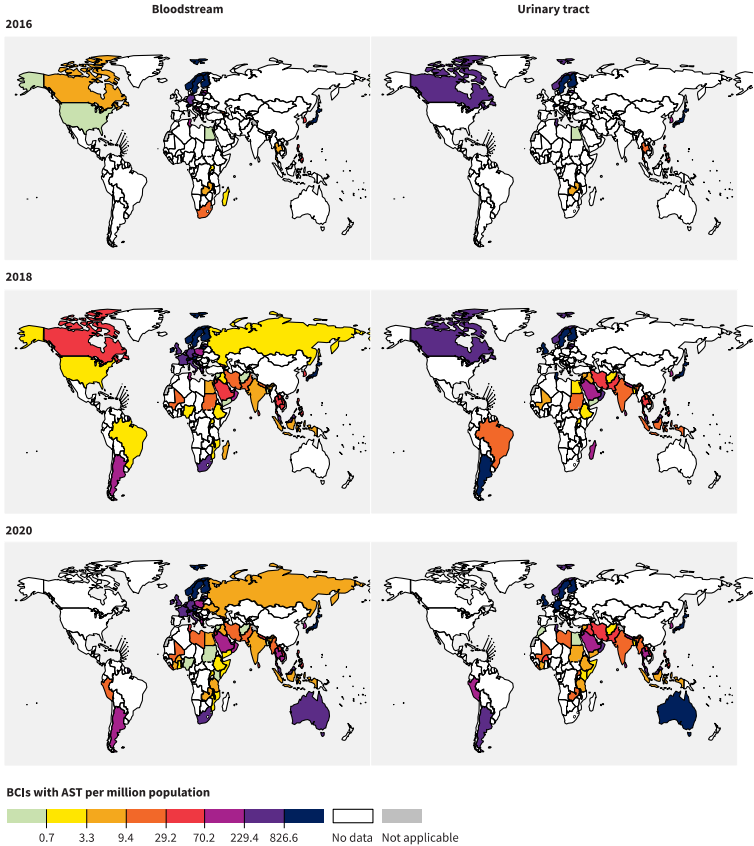
Fig. 3.2. CTAs enrolled in GLASS-AMR that reported 2016-2020 bacterial identification results and/or AST results for bacteriologically confirmed infectious syndromes under surveillance in 2017-2021 data calls



No. Participating countries

<https://iris.who.int/bitstream/handle/10665/364996/9789240062702-eng.pdf?sequence=1>

Fig. 3.4. BCIs with AST results reported to GLASS-AMR per one million population for selected infectious syndromes under surveillance (2016, 2018, 2020)



Note: In 2020, 16 CTAs reported bloodstream BCIs and 14 reported urinary tract BCIs with AST results for the first time. Five CTAs reported bloodstream BCIs in earlier years, but not in 2020, including Brazil, Canada and the United States of America, and nine reported urinary tract BCIs in earlier years, but not in 2020, including Brazil and Canada. Most CTAs reporting higher numbers of bloodstream BCIs with AST results per million population in 2020 (that is, ≥ 70.2) were from the European region (56%), whereas most CTAs reporting higher numbers of urinary tract BCIs with AST results were from the Eastern Mediterranean region (36%). Maps for gastrointestinal and gonorrhoea infectious syndromes are available as part of the web-based expanded content.

No data from Canada 2020

Comment la RAM se comporte-t-elle au Québec par rapport au reste du monde?

Agents: E. coli et K.pneumoniae
Source: bactériémies

- A. Bien mieux que la moyenne
- B. Un peu mieux que la moyenne
- C. Dans la moyenne
- D. Un peu en dessous de la moyenne
- E. Bien en-dessous de la moyenne

slido

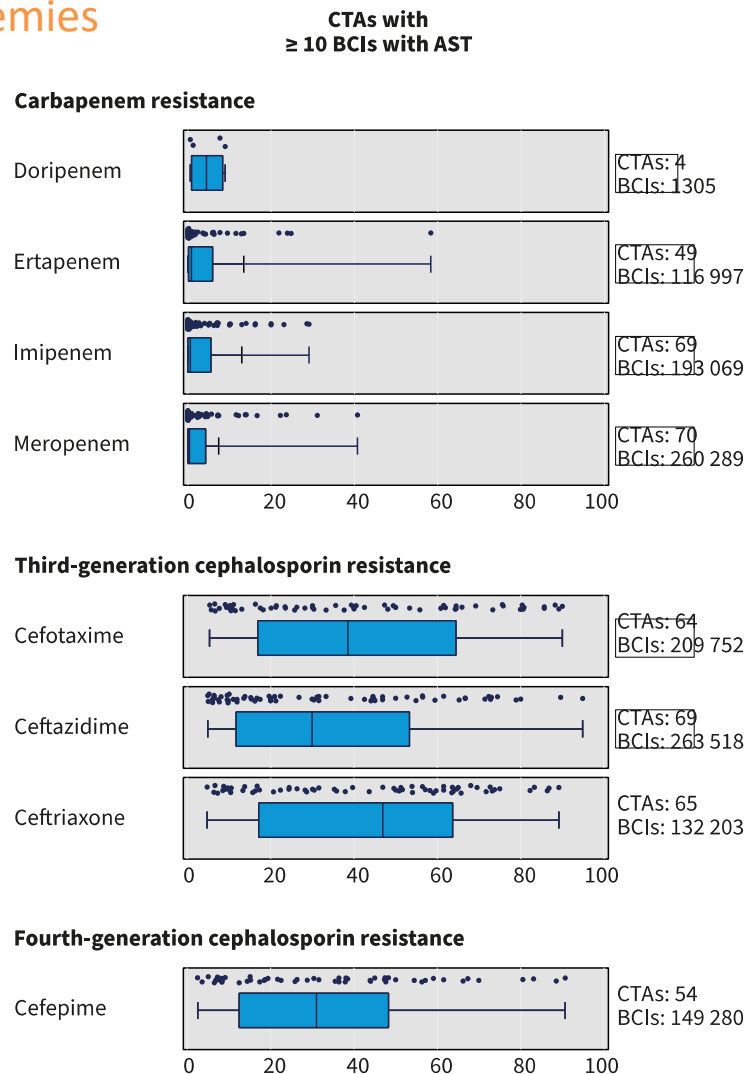
Please download and install the Slido app on all computers you use



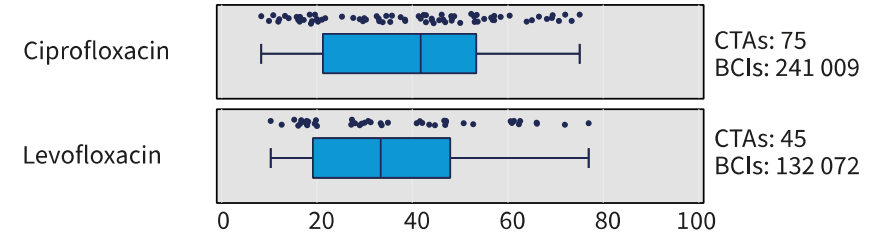
Comment la RAM se comporte-t-elle au Québec par rapport au reste du monde? (bactériémies à E. coli et K.pneumoniae)

i Start presenting to display the poll results on this slide.

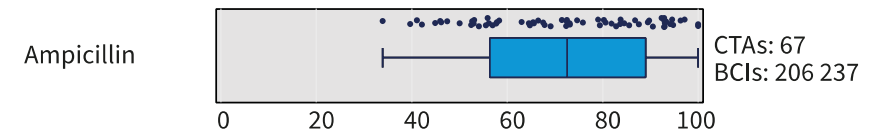
E.Coli bactériémies



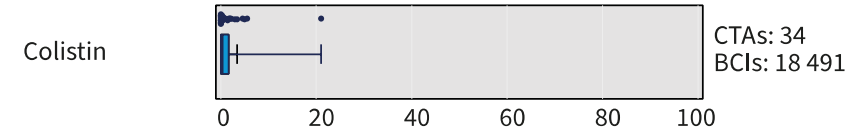
Fluoroquinolone resistance



Penicillin resistance



Polymyxin resistance



Sulfonamide and trimethoprim resistance

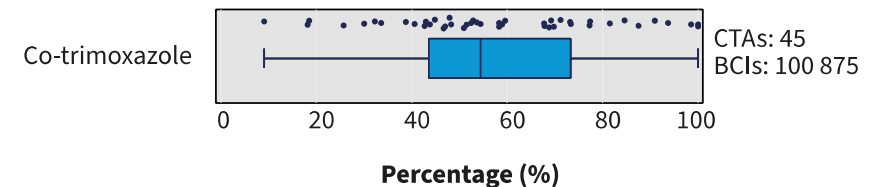
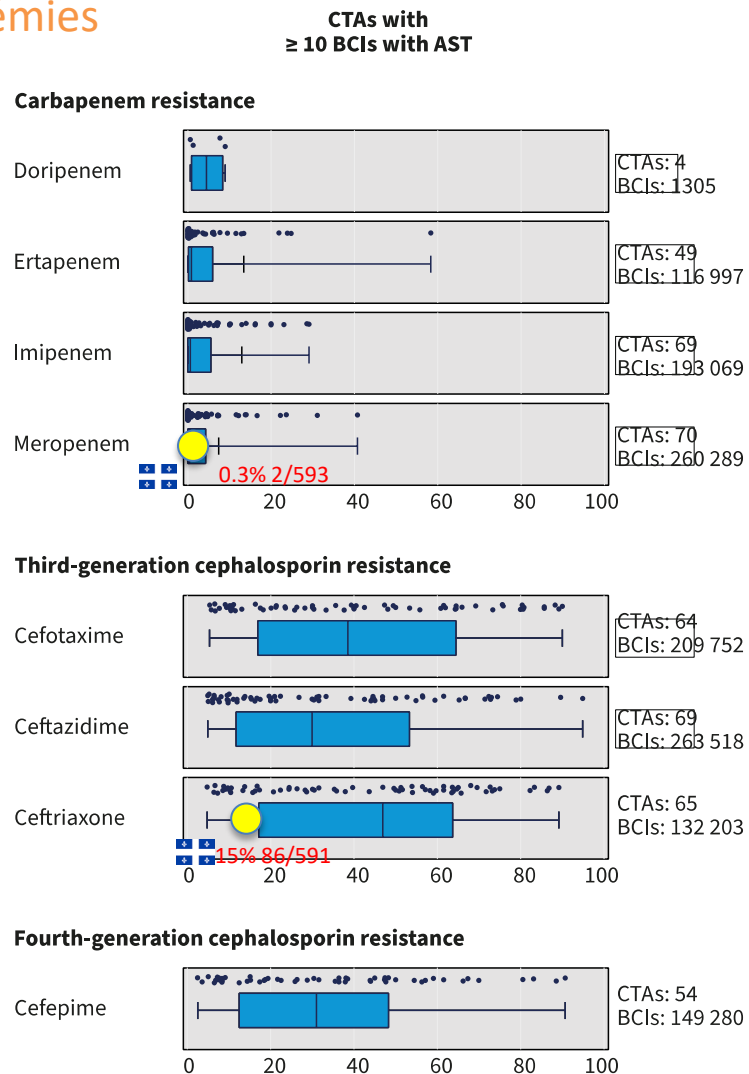
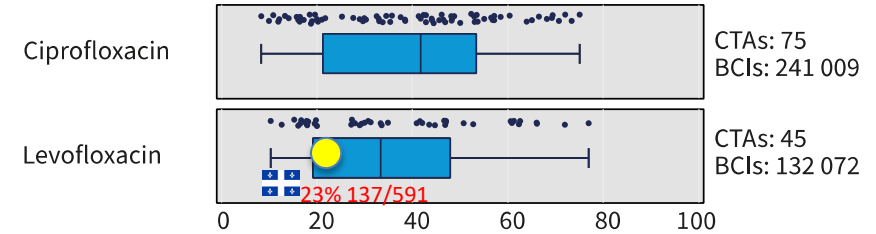


Fig. 3.7b. Percentage resistance to antimicrobials under surveillance in *E. coli* in all CTAs reporting ≥ 10 *E. coli* bloodstream infections with AST results compared to CTAs where the reported number per million population was above the 75th percentile in 2020

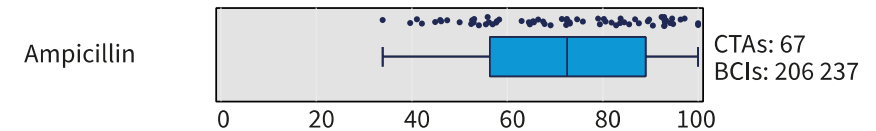
E.Coli bactériémies



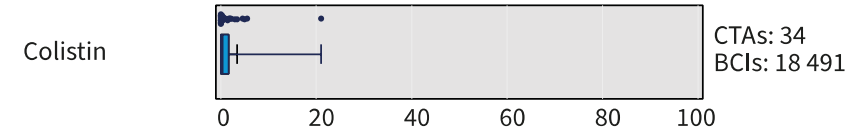
Fluoroquinolone resistance



Penicillin resistance



Polymyxin resistance



Sulfonamide and trimethoprim resistance

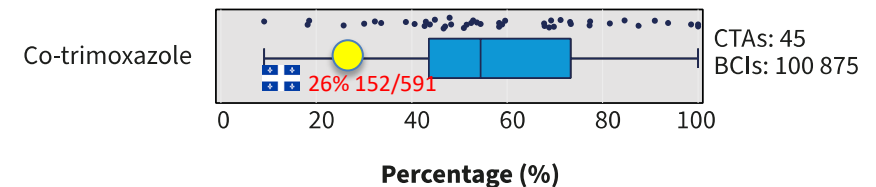


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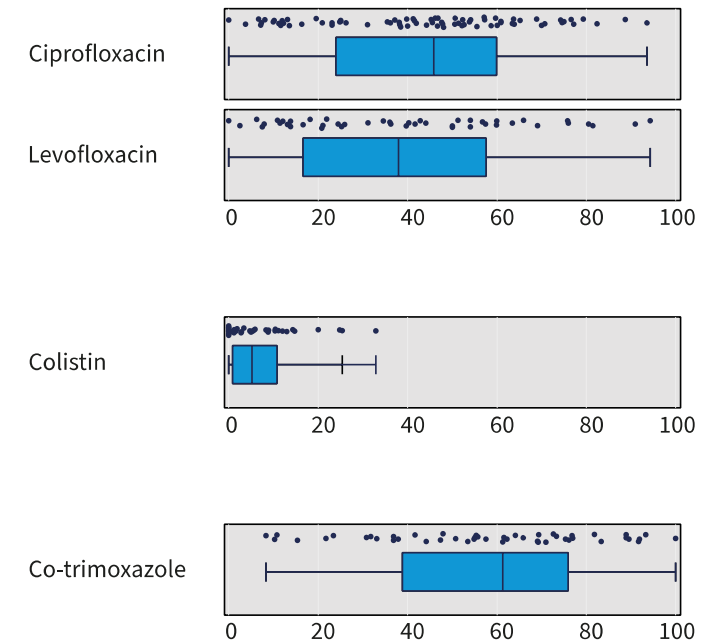
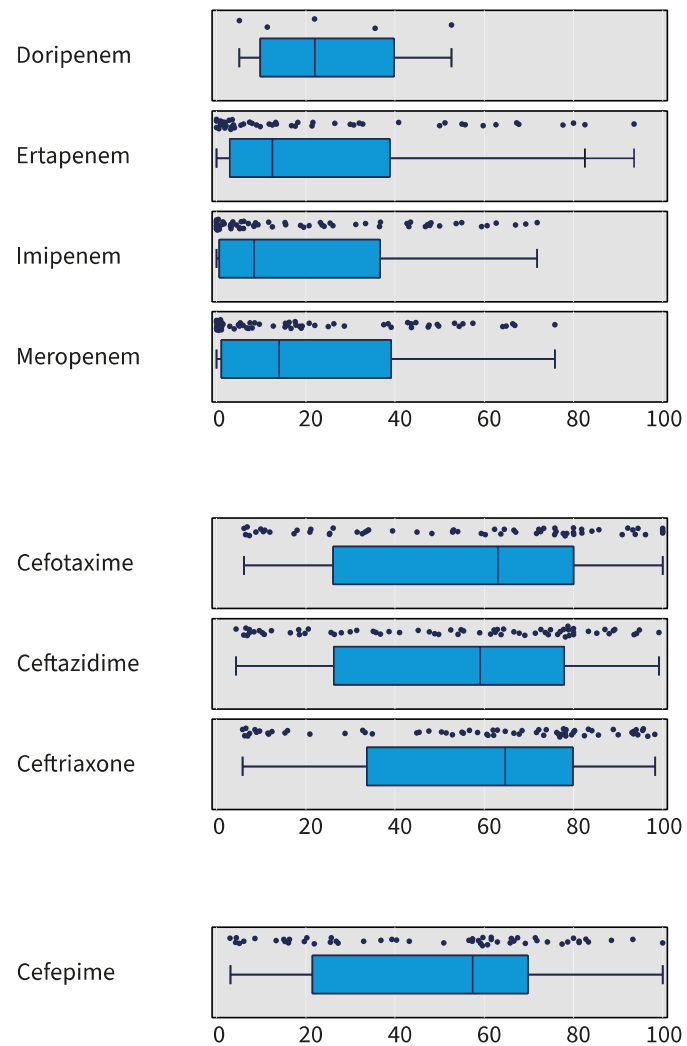


Fig. 3.7c. Percentage resistance to antimicrobials under surveillance in *K. pneumoniae* in all CTAs reporting ≥ 10 *K. pneumoniae* bloodstream infections with AST results compared to CTAs where the reported number per million population was above the 75th percentile in 2020

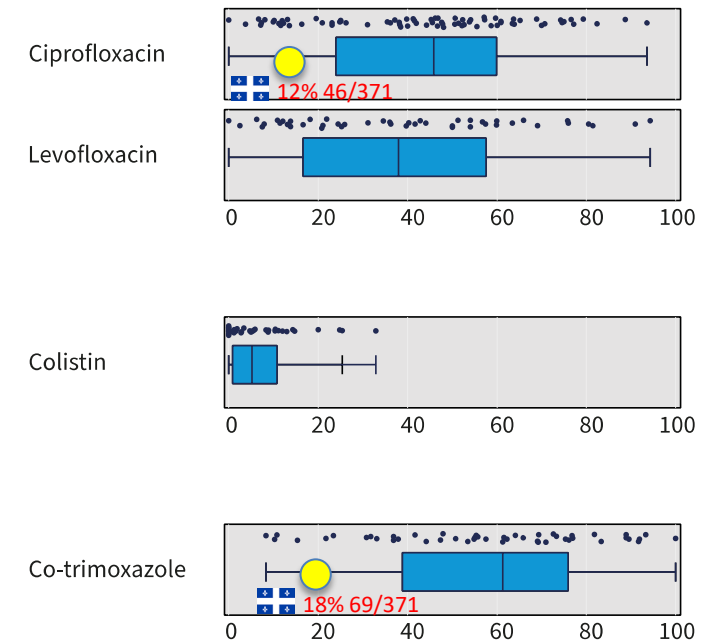
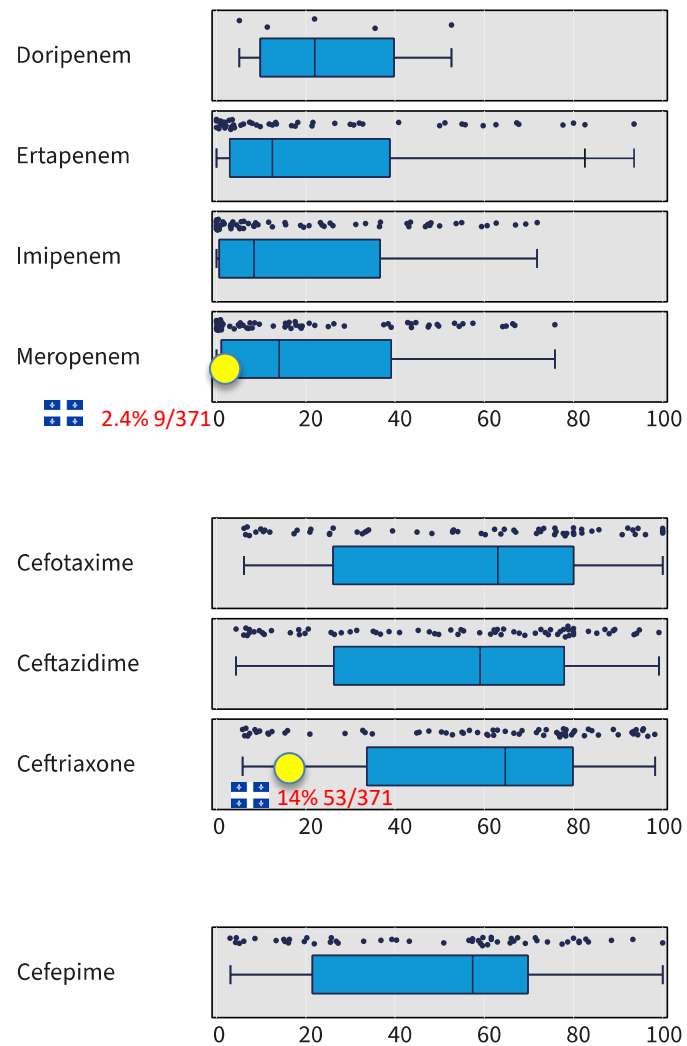


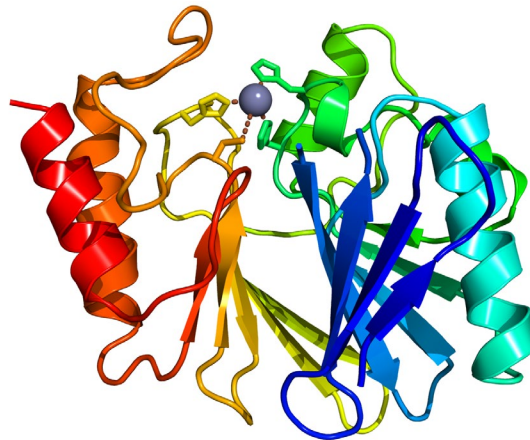
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Global burden of bacterial antimicrobial resistance 1990–2021: a systematic analysis with forecasts to 2050

GBD 2021 Antimicrobial Resistance Collaborators*

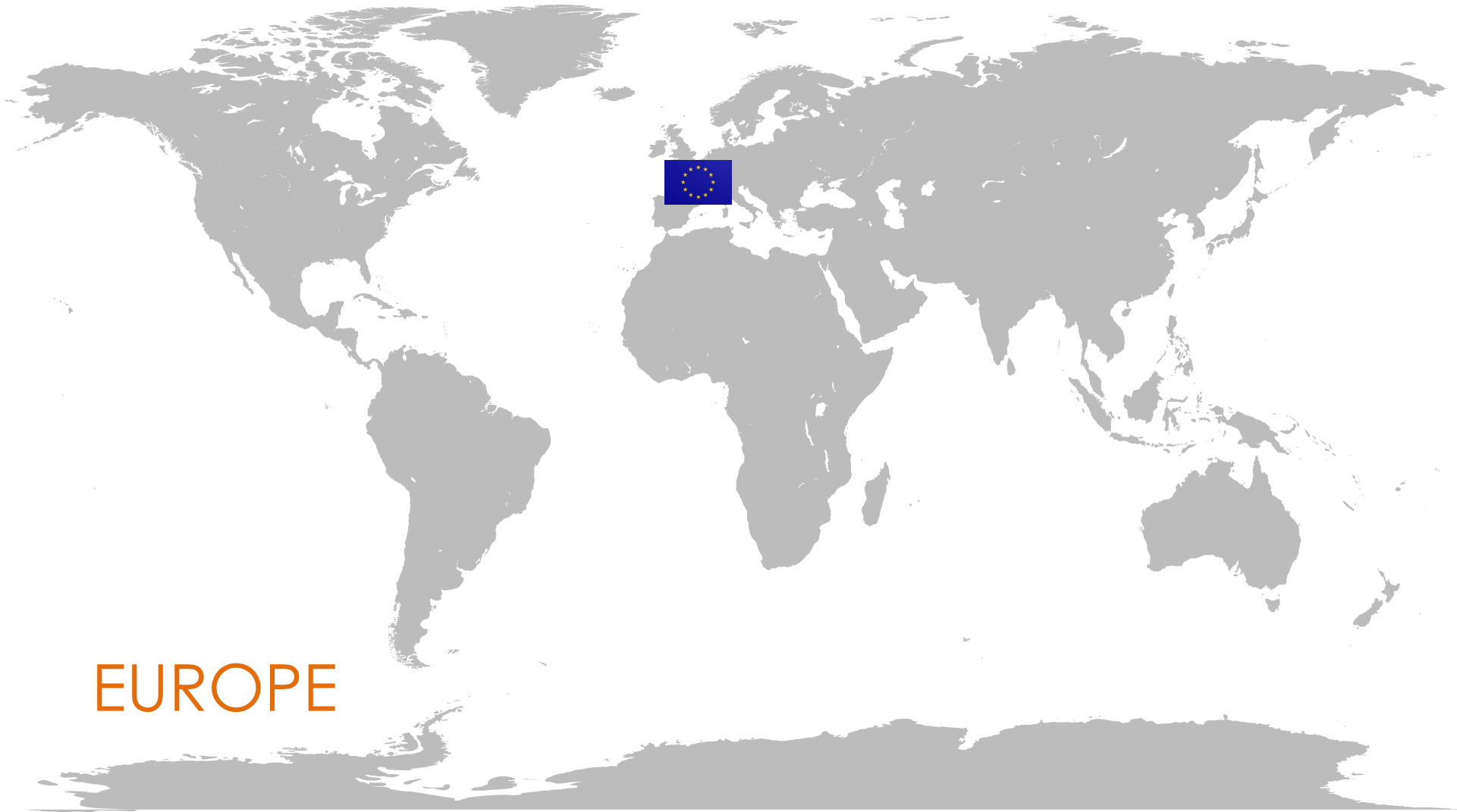


Les bactéries GN résistantes aux carbapénèmes est la forme de RAM qui devrait augmenter le plus

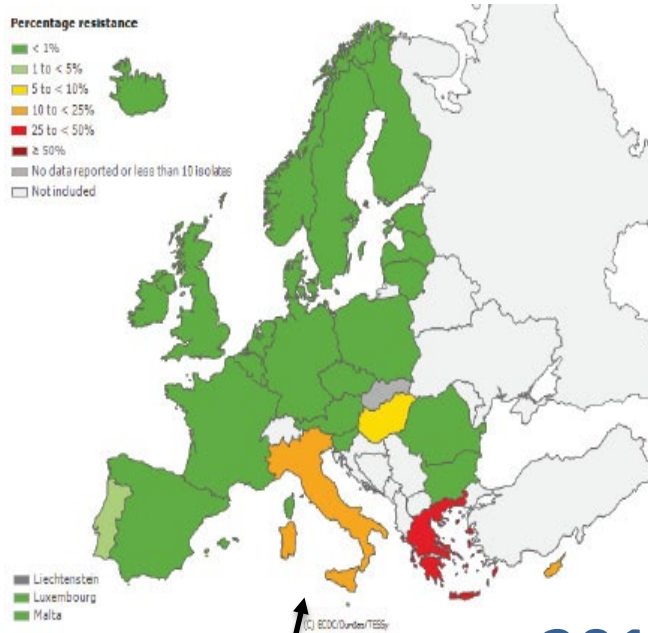


Metallo-B-lactamase

	1990	2021
Associated deaths	619 000	1 030 000
Attributable deaths	127 000	216 000



Klebsiella pneumoniae : pourcentage d'isolats invasifs résistants aux carbapénèmes.



2010

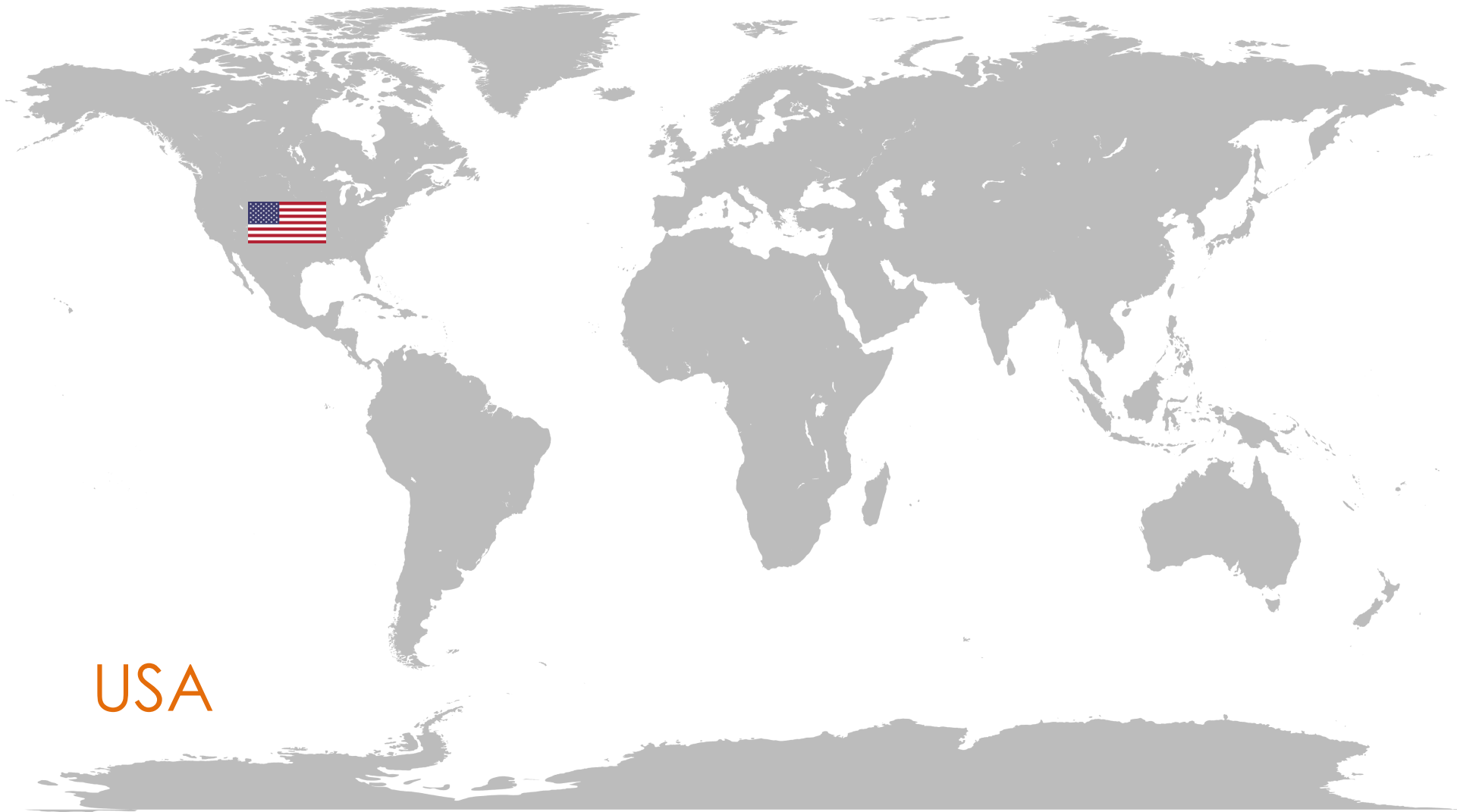


2011

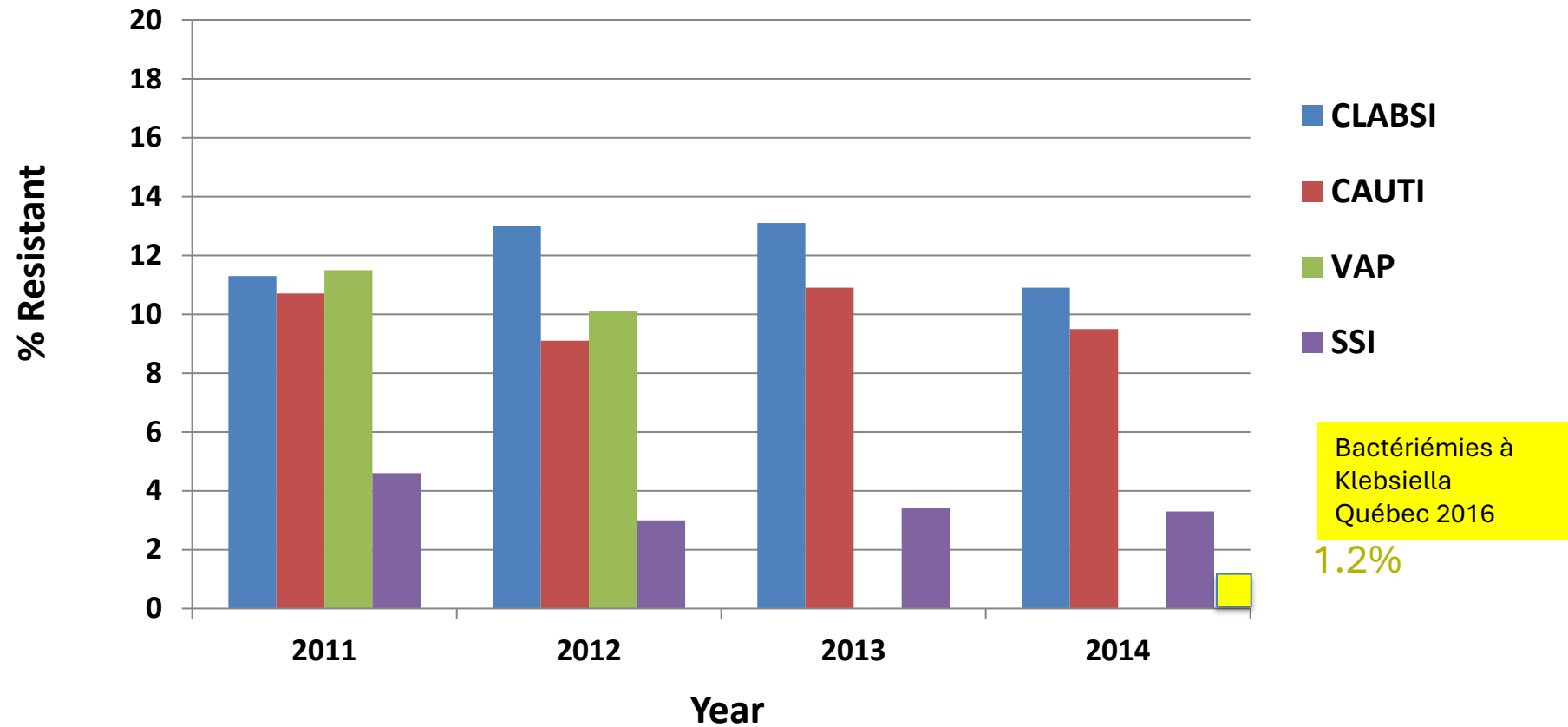


2014

ITALIE (2006): 1.3%



% carbapenem-resistant *Klebsiella* species (CRE)



Source: A. Lee U. Sydney

Weiner et al. Infect Control Hosp Epidemiol 2016; 37: 1288-1301

Transmission d'EPC de la Grèce à d'autres pays européens 2007-2010

Country	Year	Total Number of Patients	Origin of Patients	Number of Secondary Cases	Probability of the Greek Origin	References	Mechanisms of Resistance
Belgium	2009	3	3 patients transferred from Greek hospitals	0	Confirmed	Bogaerts et al. 2010 [19]	<i>blaKPC-2</i>
Denmark	2009	2	2 patients transferred from Greek hospitals	0	Confirmed	Hammerum et al. 2010 [20]	<i>blaKPC-2</i>
Finland	2009	1	1 patient transferred from Crete	0	Confirmed	Osterblad et al. 2010 [21]	<i>blaKPC-2</i>
France	No data	8	1 patient transferred from Crete	7	Confirmed	Naas et al. 2010 [22]	<i>blaKPC-2</i>
France	2007	1	1 patient transferred from Crete	0	Confirmed	Cuzon et al. 2008 [23]	<i>blaKPC-2</i>
France	2009	1	1 patient transferred from Greek hospital	0	Confirmed	Barbier et al. 2010 [24]	<i>blaKPC-2</i>
France	2009	4	1 patient transferred from Greek hospital	3	Confirmed	Kassis-Chikhani et al. 2010 [25]	<i>blaKPC-2</i>
Germany	2007-2008	9	1 patient treated in Greece	8	Hypothetical	Wendt et al. 2010 [26]	<i>blaKPC-2</i>
Hungary	2008	7	1 patient transferred from Greek hospital	6	Confirmed	Tóth et al. 2010 [27]	<i>blaKPC-2</i>
Norway	2007	6	4 patients transferred from Greek hospitals	2	Confirmed	Samuelson et al. 2009 [28]	<i>blaKPC-2</i>
Sweden	No data	1	1 patient transferred from Greek hospital	0	Confirmed	Mark Wisell et al. 2007	<i>blaKPC-2</i>
The Netherlands	No data	14	1 patient transferred from Greek hospital	13	Confirmed	et al. 2010 [30]	<i>blaKPC-2</i>



2011



CNISP CPO Surveillance Update

Mise à jour de la surveillance OPC du PCSIN

CNISP annual meeting/Réunion annuelle du PCSIN

October 2023/octobre 2023



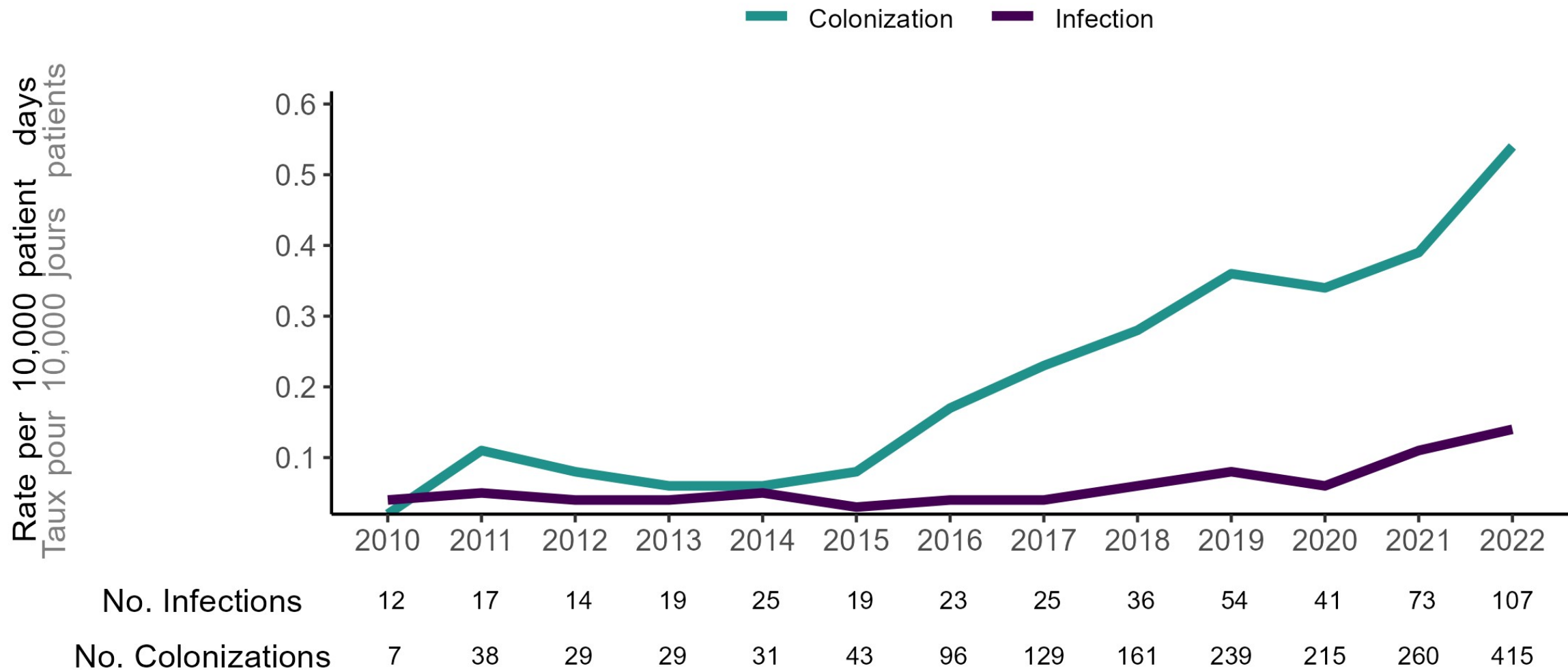
Public Health
Agency of Canada

Agence de la santé
publique du Canada

CPE infection and colonization rates, 2010-2022

Taux d'infection et colonisation de l'EPC, 2010-2022

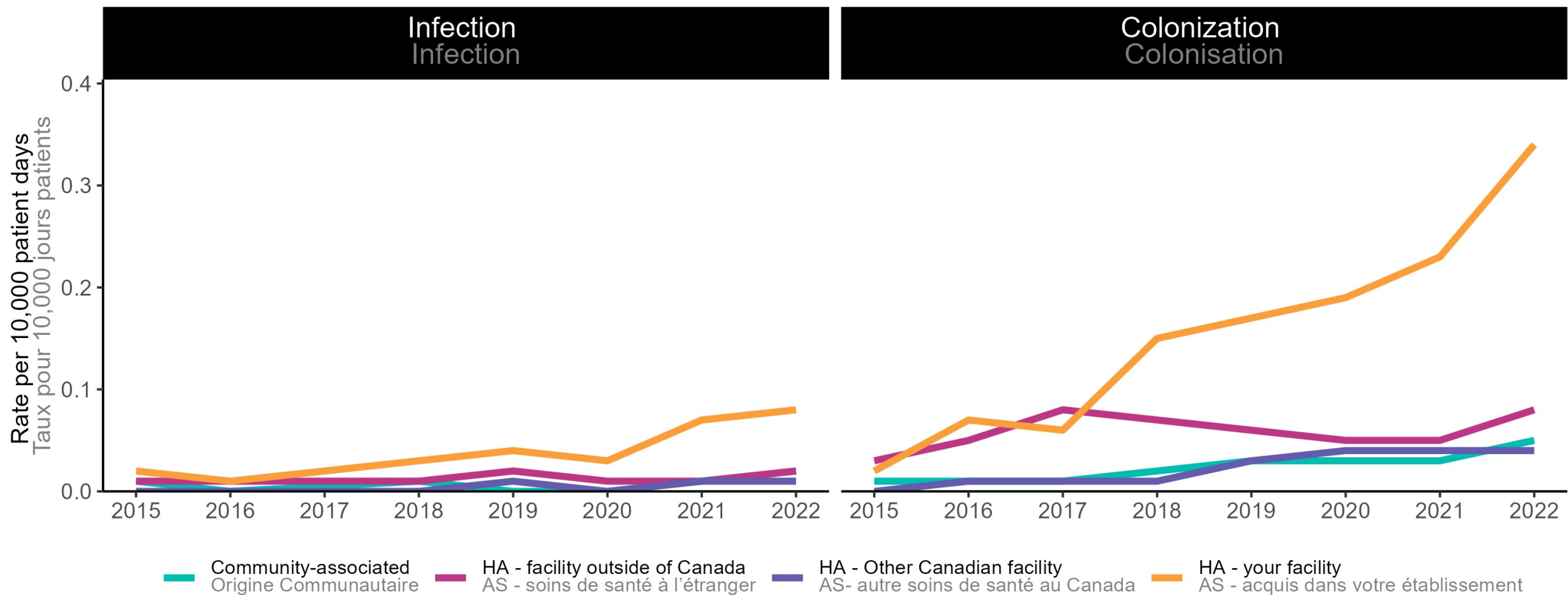
CPE



CPE rates by source of acquisition, 2015-2022

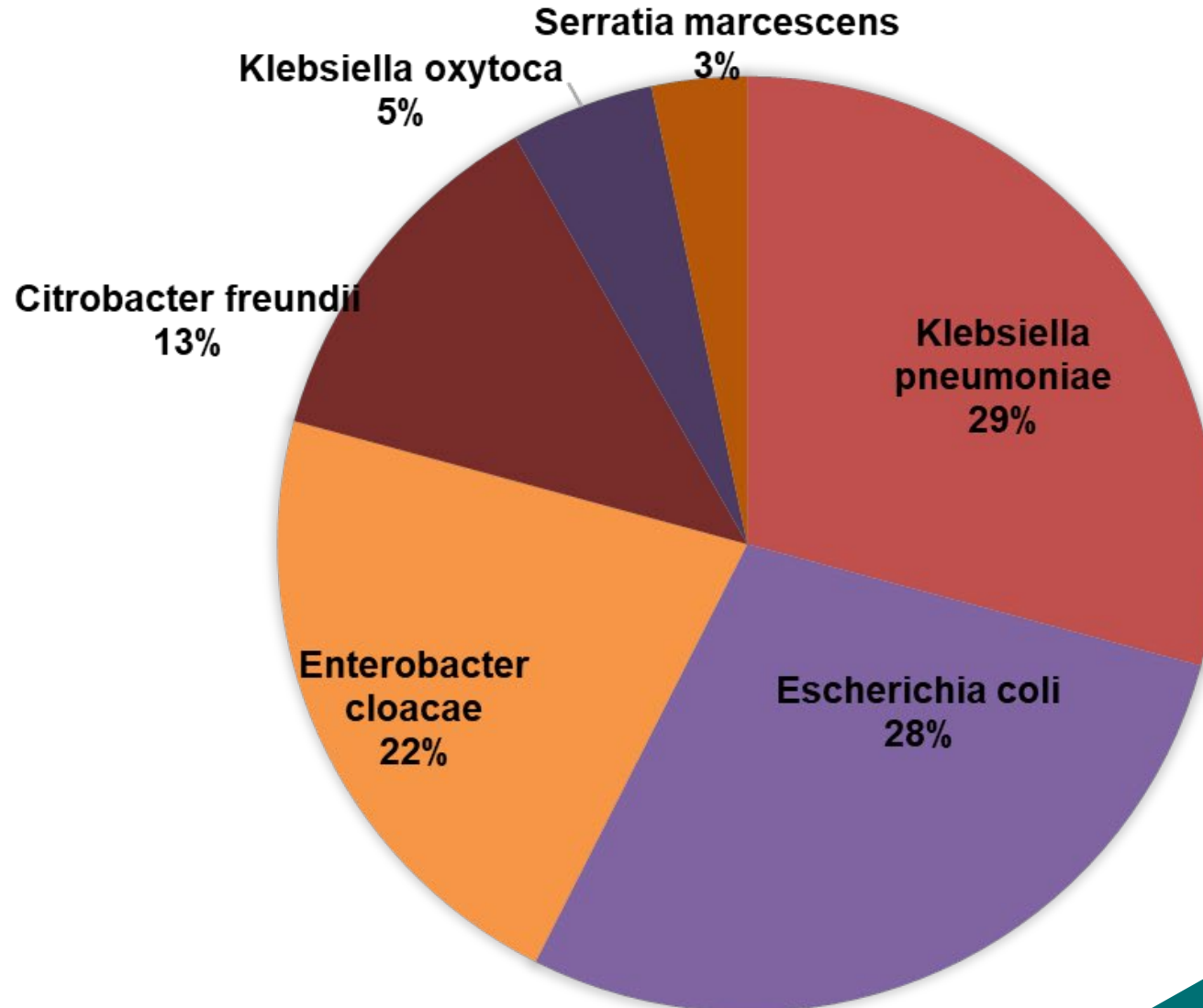
Taux d'EPC selon la source d'acquisition, 2015-2022

CPE



Proportion of carbapenemase producing Enterobacteriaceae (CPE)

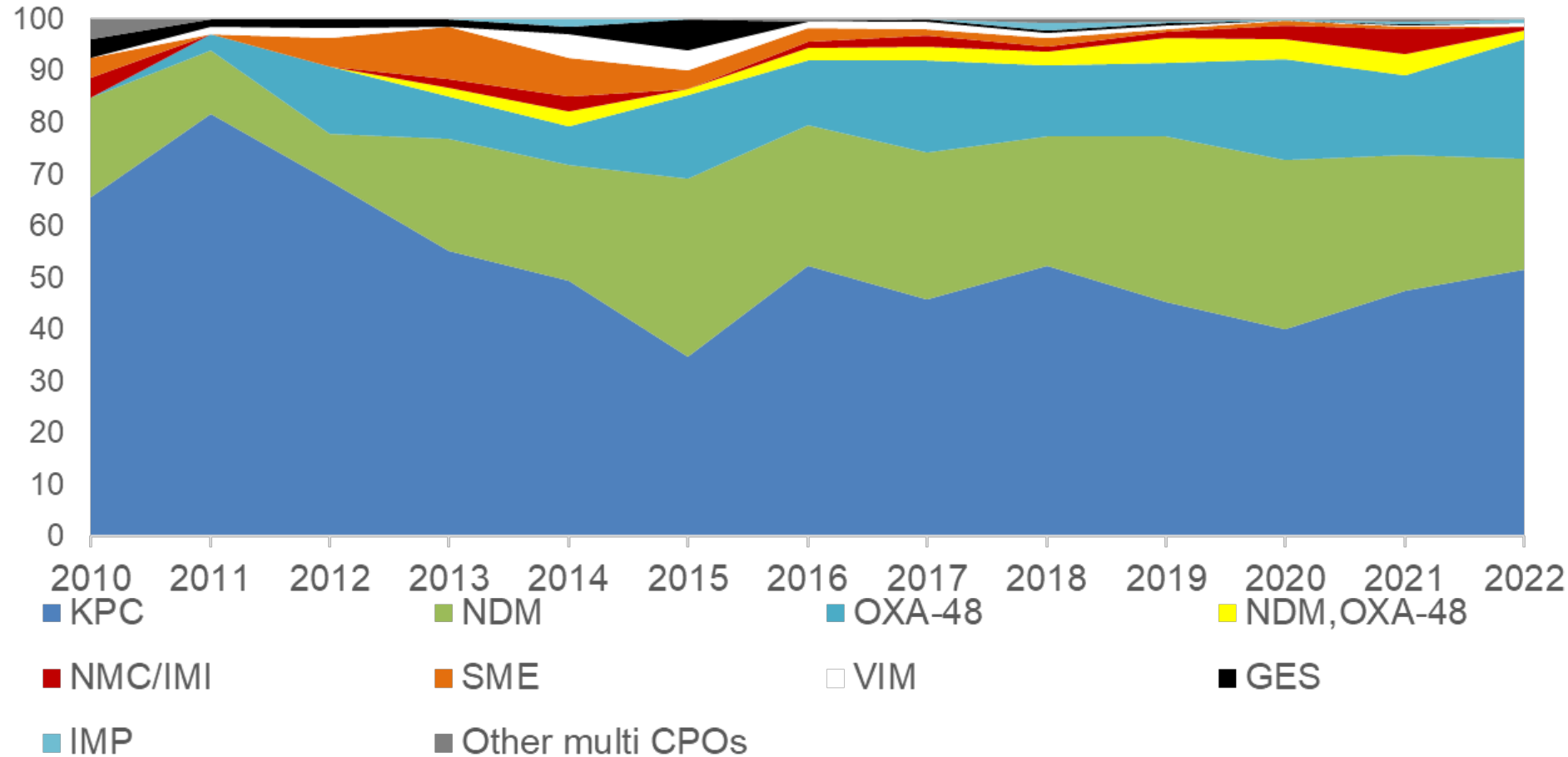
NML



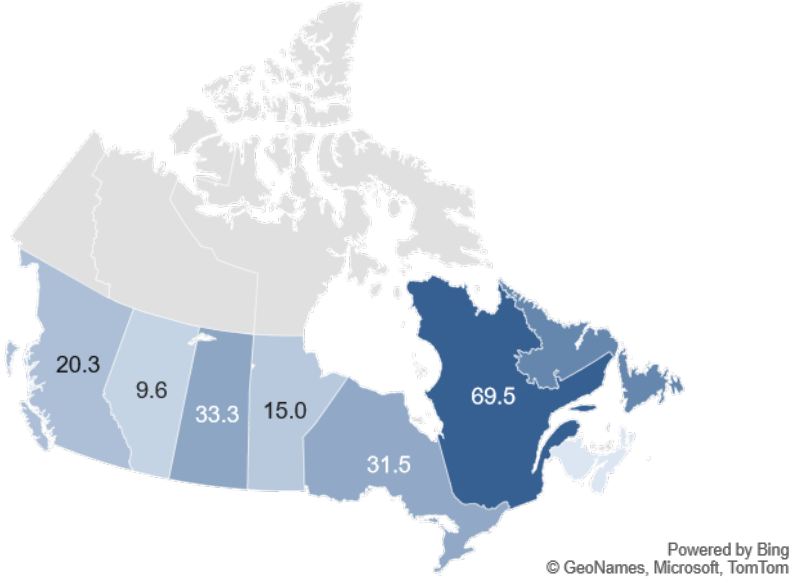
Proportion CPE by year

NML

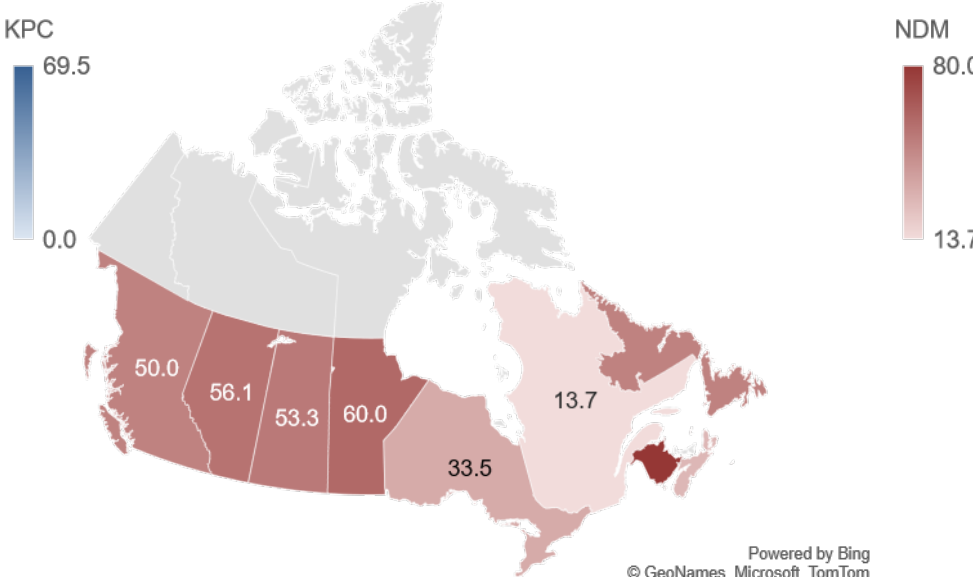
91% of CPE
harbour NDM,
KPC or OXA-48



Proportion of KPC within each province

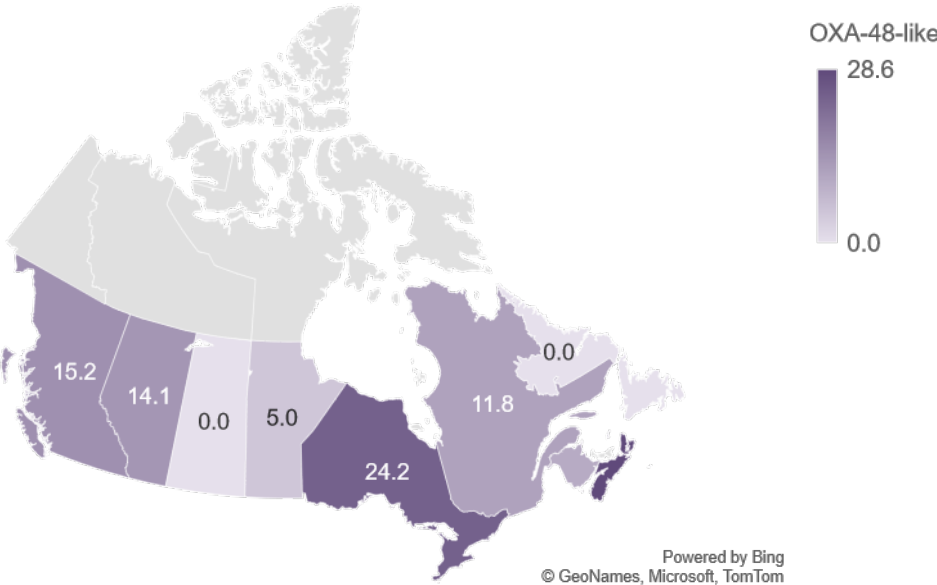


Proportion of NDM within each province



NML

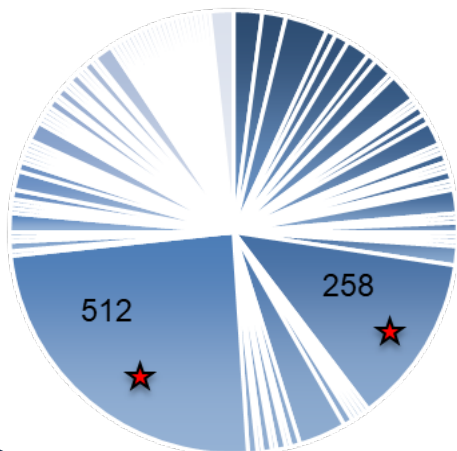
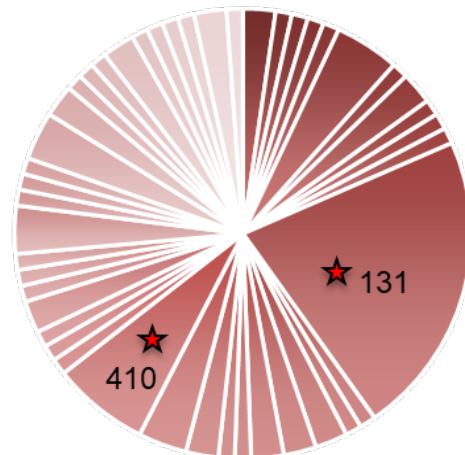
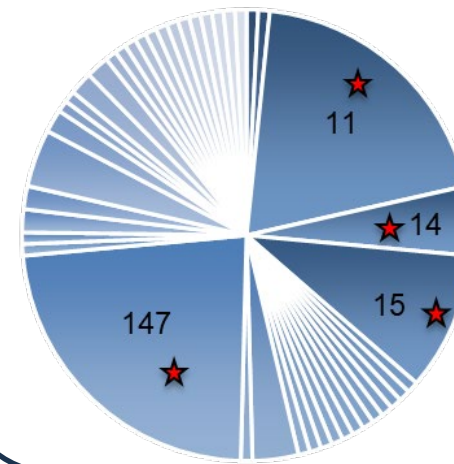
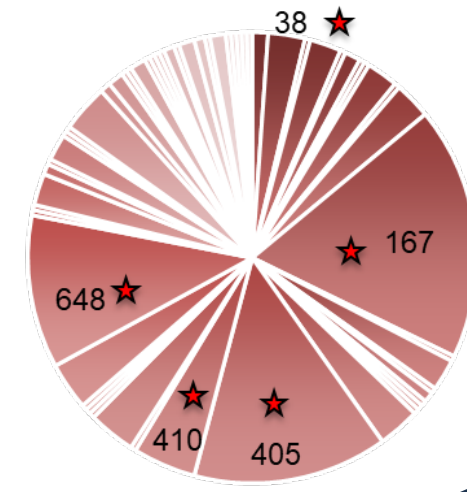
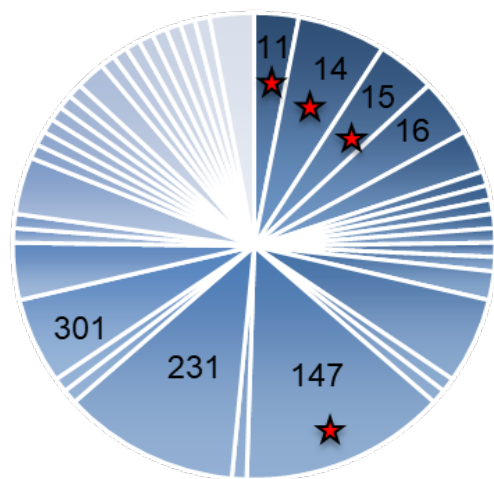
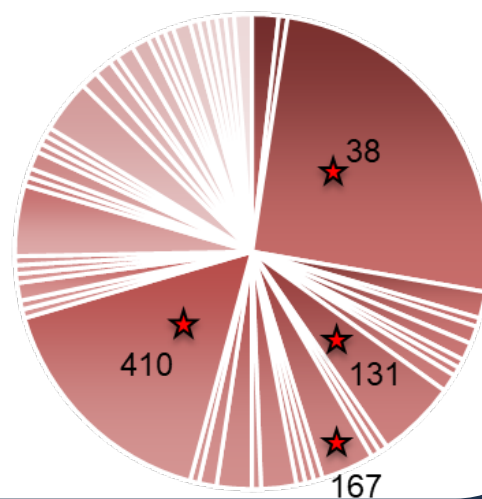
Proportion of OXA-48-like within each province



NDM-BC, AB, SK, MB
KPC-QC
OXA-48-ON

Typing the top CPOs-*K. pneumoniae* and *E.coli*

NML

KPC-*K.pneumoniae*KPC-*E.coli*NDM-*K.pneumoniae*NDM-*E.coli*OXA-48-like-*K.pneumoniae*OXA-48-like-*E.coli*

★ High risk clone

QUÉBEC



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Le réviseur a été convié à apporter des commentaires sur la version préfinale de ce document et en conséquence, n'en ont pas révisé ni endossé le contenu final.

Les auteur(-trice)s ainsi que les membres du comité scientifique et les réviseurs ont dûment rempli leurs déclarations d'intérêts et aucune situation à risque de conflits d'intérêts réels, apparents ou potentiels n'a été relevée.

FINANCEMENT

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MISE EN PAGE

Marie-Amélie Bras, agente administrative
Direction des risques biologiques

REMERCIEMENTS

Nous tenons à remercier toutes les équipes de prévention et contrôle des infections qui participent à la surveillance des infections nosocomiales au Québec.

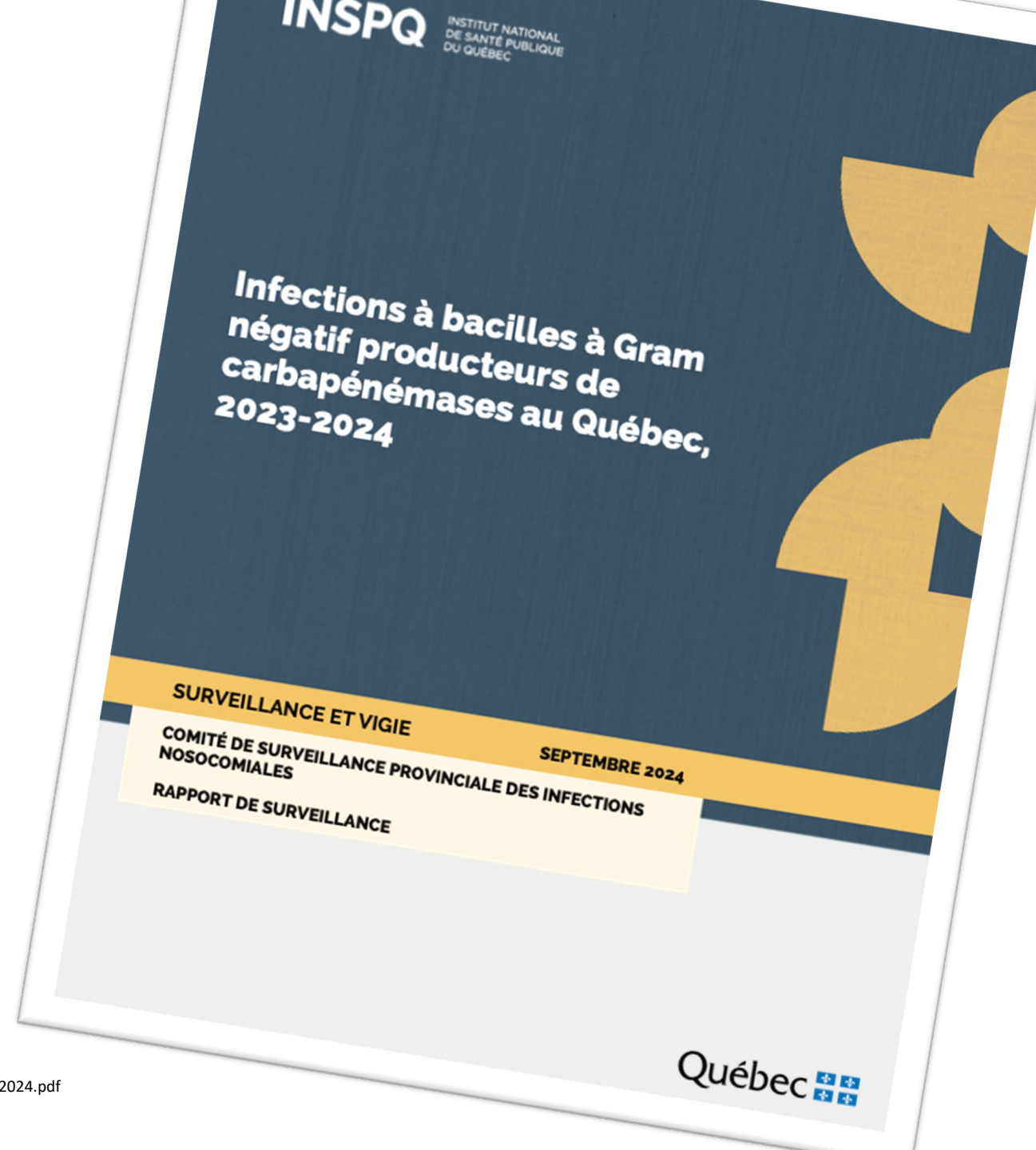


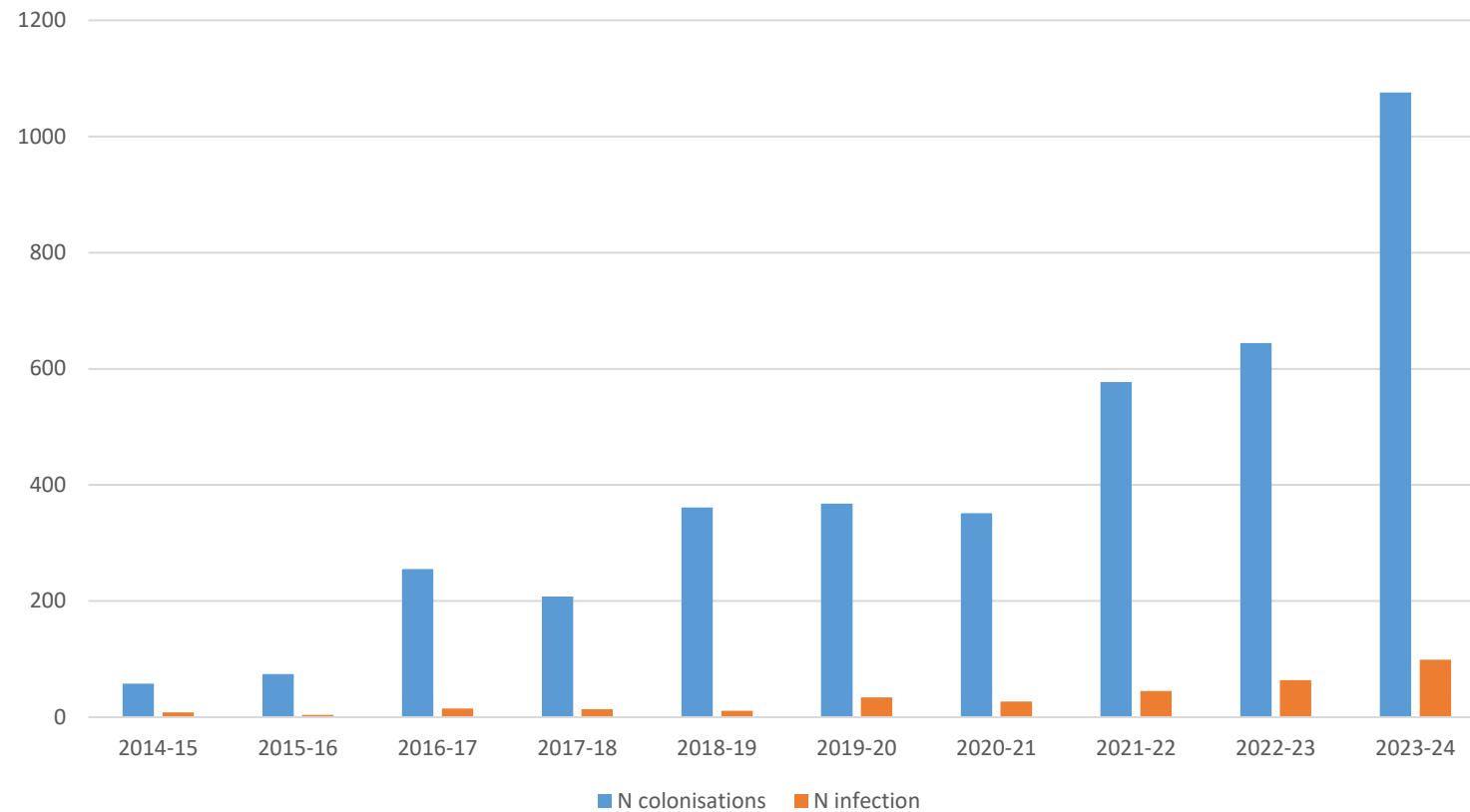
Tableau 1 Sommaire de la participation, des cas, des dénominateurs et des taux pour la surveillance des infections à BGNPC, 2019-2020 à 2023-2024

	2019-2020	2020-2021	2021-2022	2022-2023	2023-2024
Installations participantes (N)	88	84	83	86	88
Admissions (N)	735 713	596 209	648 874	665 604	688 231
Jours-présence (N)	4 965 755	4 183 386	4 409 071	4 666 352	4 828 333
Infections à BGNPC					
Infections nosocomiales à BGNPC (cat. 1a et 1b) (N)	13	16	29	30	54
Usagers infectés (cat. 1a et 1b) (N)	12	15	29	28	49
Infections toutes catégories à BGNPC (cat. 1a, 1b, 1c, 1d, 1e, 2, 3 et 4) (N)	34	27	45	64	99
Taux d'incidence des infections nosocomiales (cat. 1a et 1b) ^A	0,03	0,04	0,07	0,06	0,11
Colonisations à BGNPC					
Colonisations nosocomiales à BGNPC (cat. 1a et 1b) (N)	368	351	577	644	1 076
Colonisations toutes catégories à BGNPC (cat. 1a, 1b, 1c, 1d, 1e, 2, 3 et 4) (N)	480	425	662	785	1 259
Taux d'acquisition des colonisations nosocomiales (cat. 1a et 1b) ^A	0,74	0,84	1,31	1,38	2,23

^A Taux par 10 000 jours-présence.

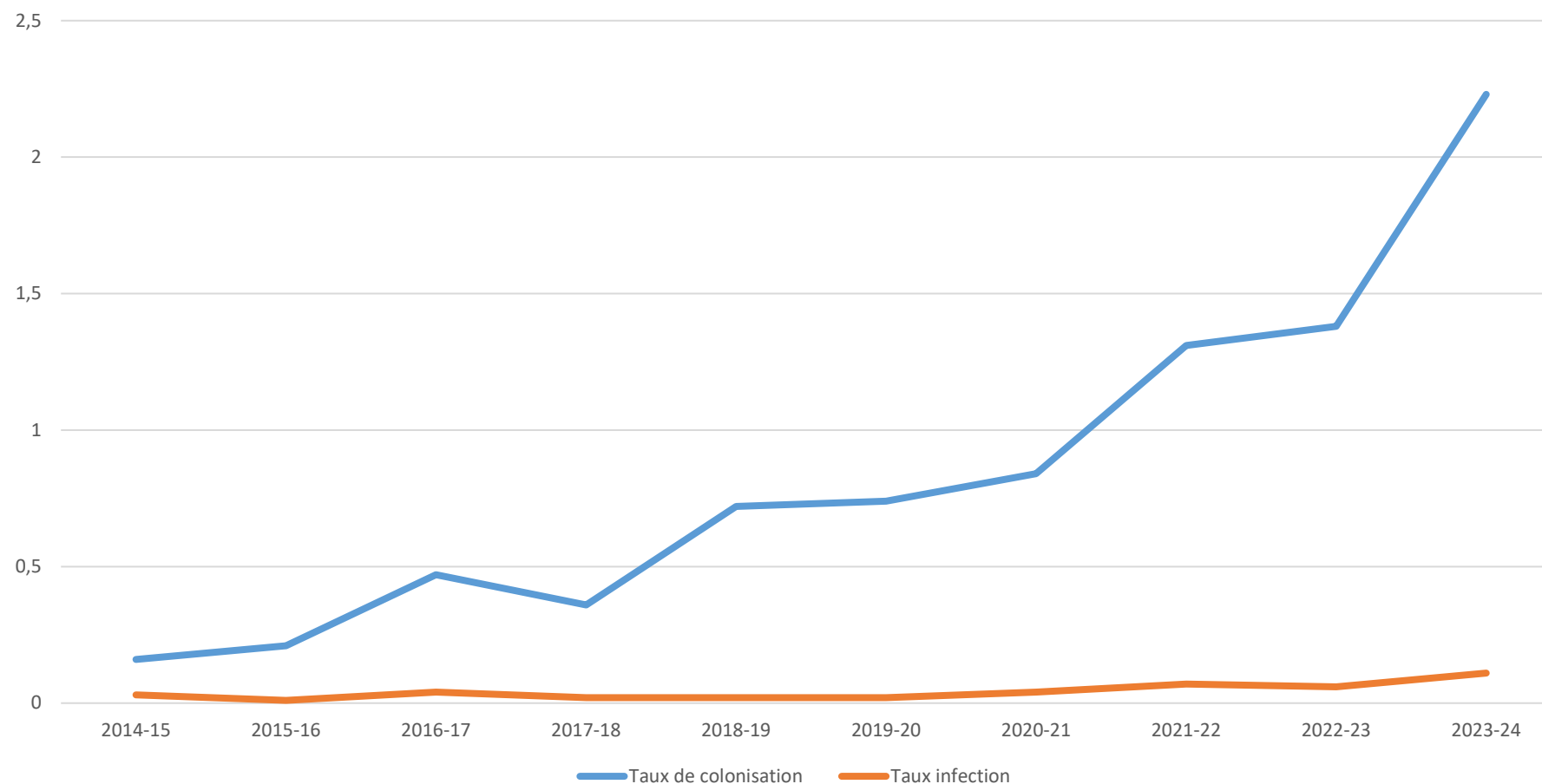
<https://www.inspq.qc.ca/sites/default/files/publications/3577-infections-bacilles-gram-negatif-carbapenemases-QC-2023-2024.pdf>

Evolution du nombre d'infections et de colonisations au Québec, 2014-2015 à 2023-2024



<https://www.inspq.qc.ca/sites/default/files/publications/3577-infections-bacilles-gram-negatif-carbapenemases-QC-2023-2024.pdf>

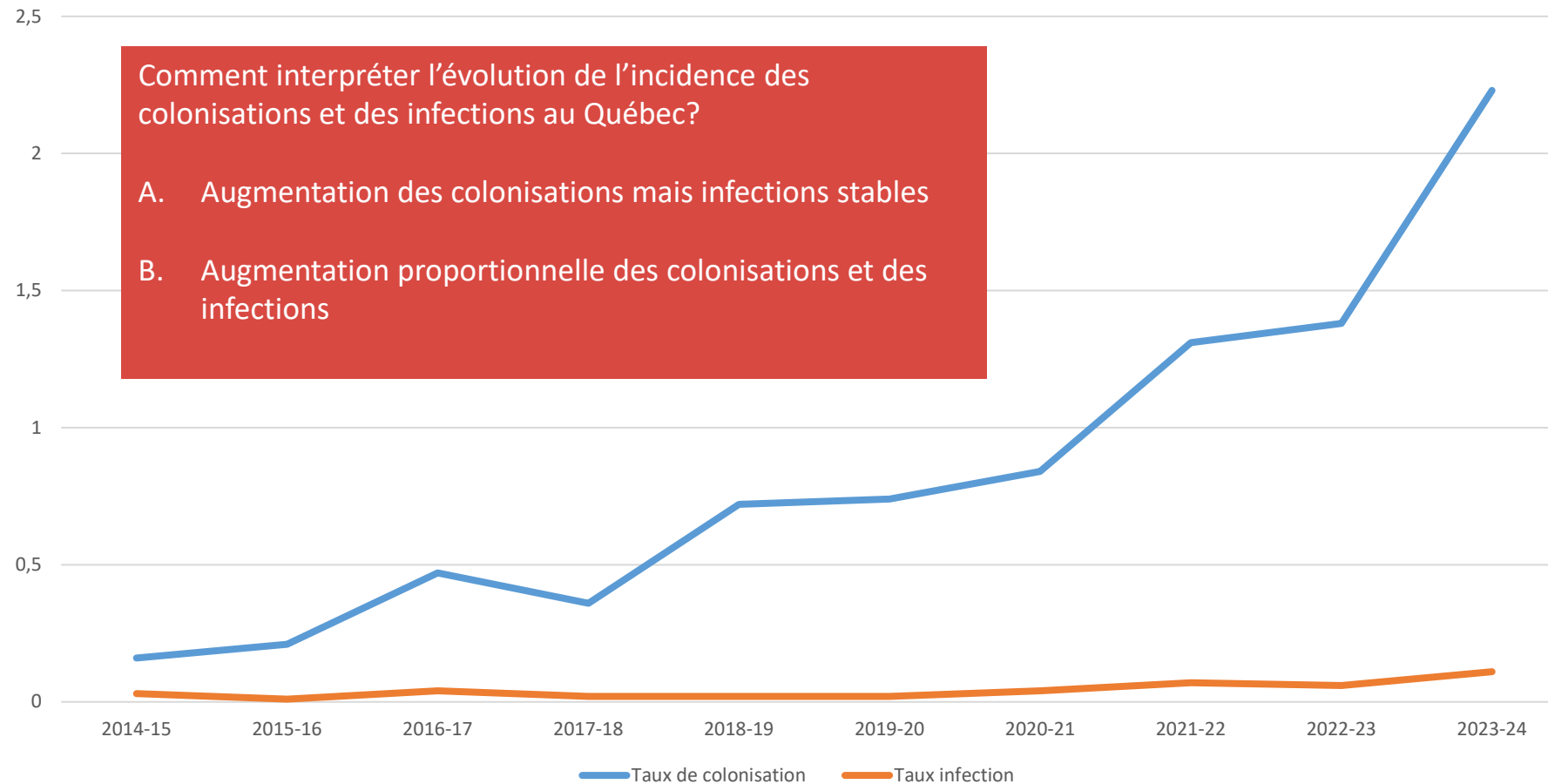
Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



- A. Augmentation des colonisations mais infections stables
- B. Augmentation des colonisations et des infections proportionnelle

<https://www.inspq.qc.ca/sites/default/files/publications/3577-infections-bacilles-gram-negatif-carbapenemases-QC-2023-2024.pdf>

Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



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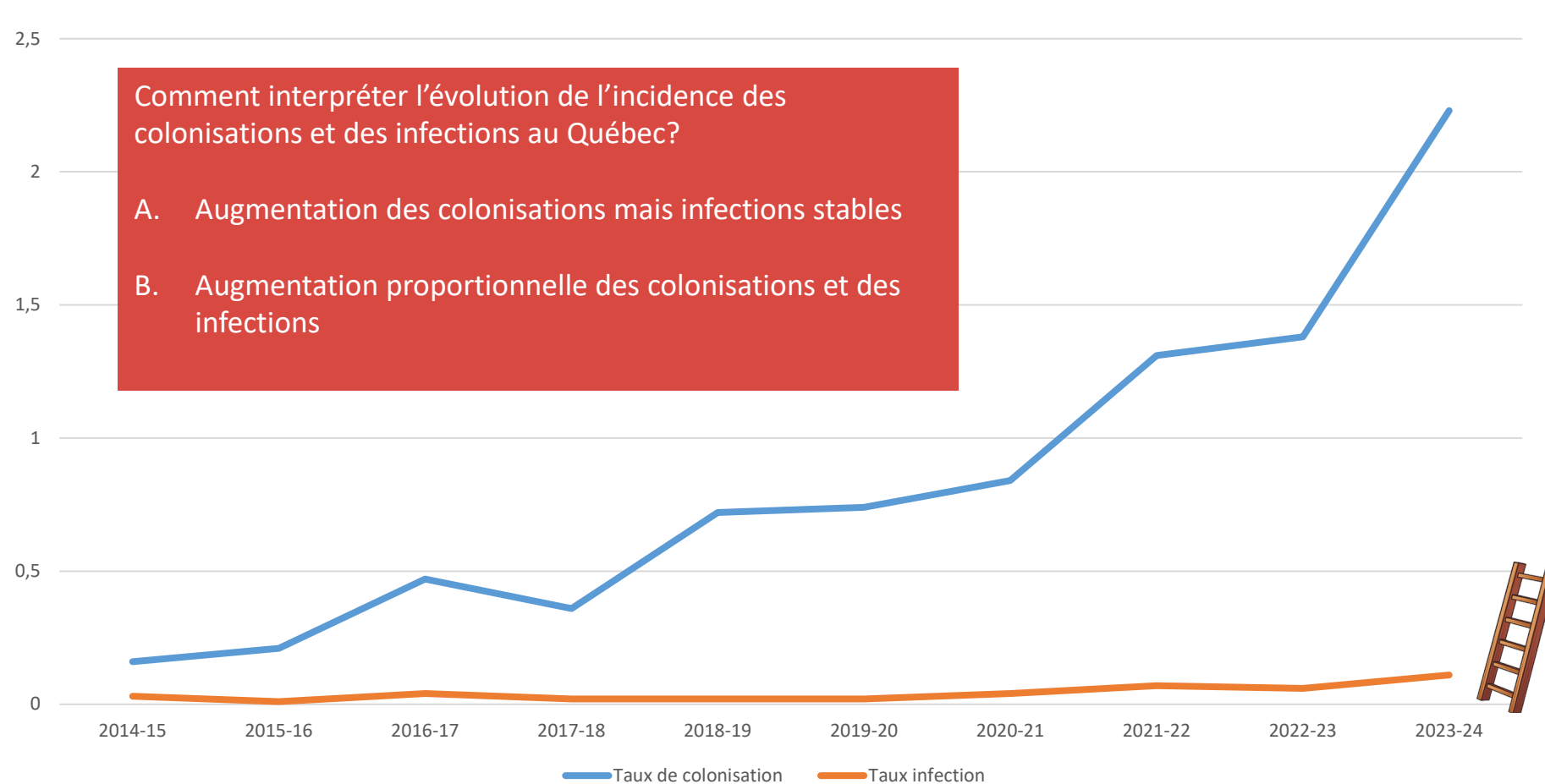
Please download and install the Slido app on all computers you use



Comment interpréter l'évolution de l'incidence des colonisations et des infections au Québec?

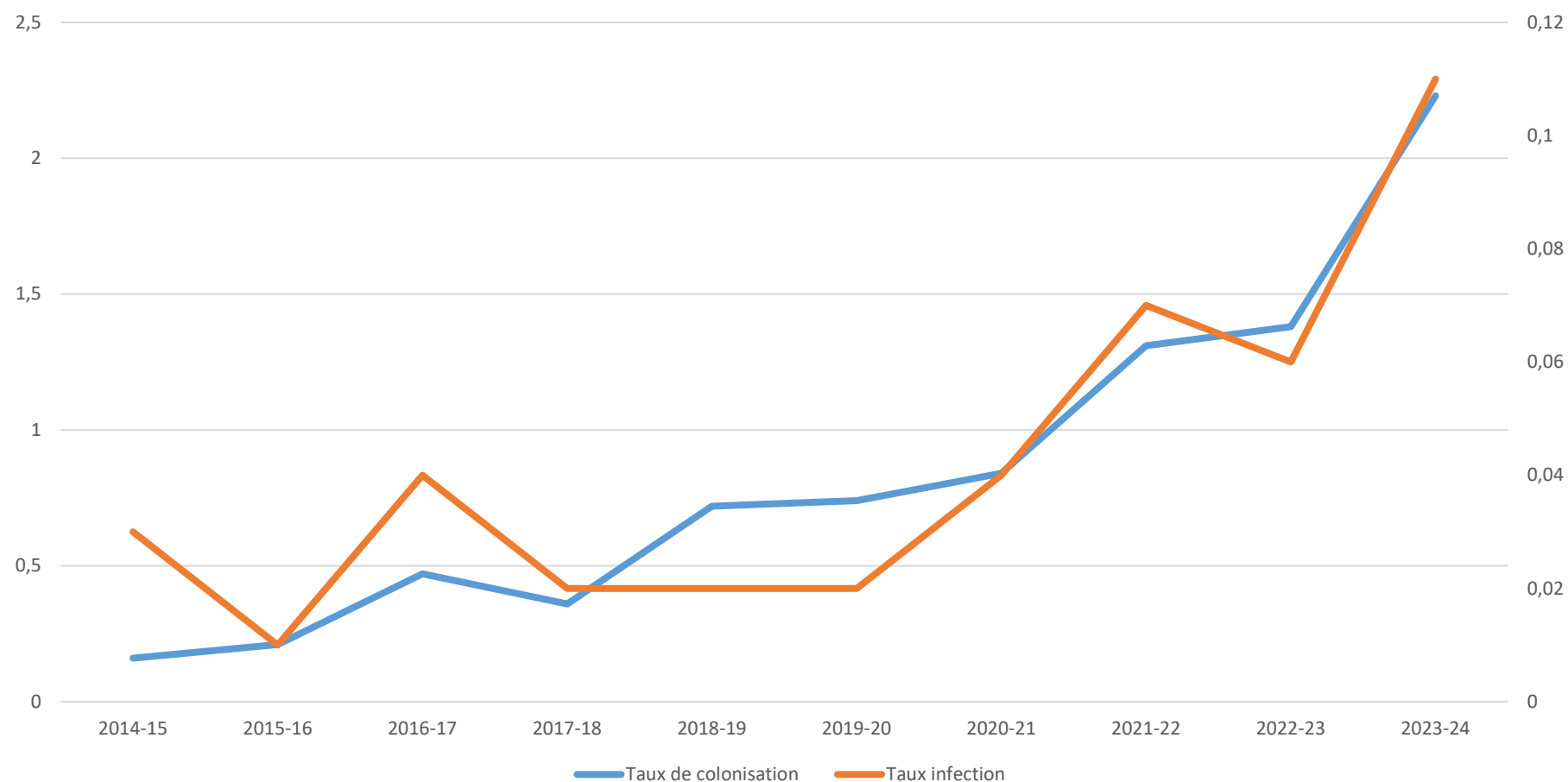
① Start presenting to display the poll results on this slide.

Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



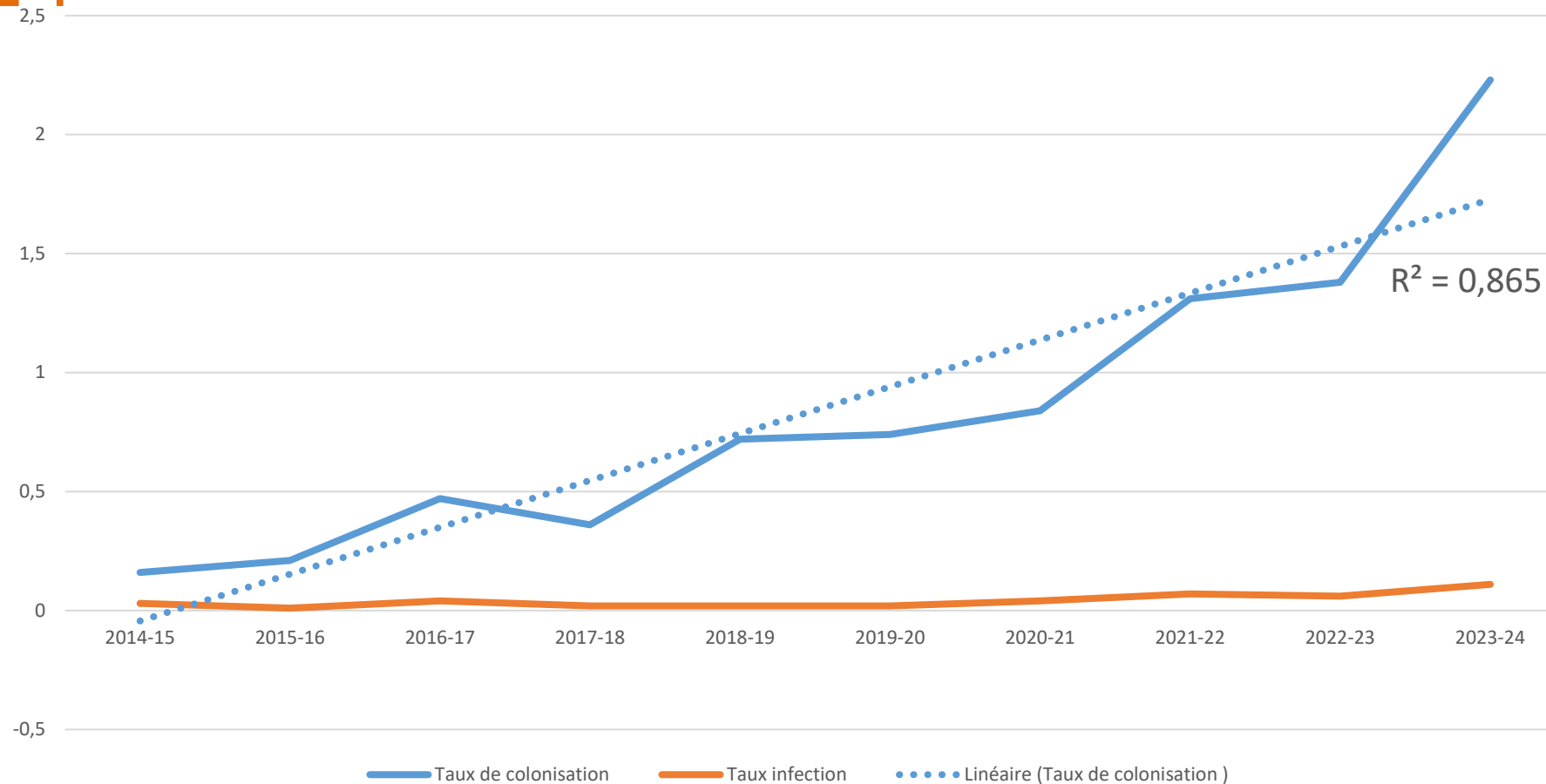
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Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



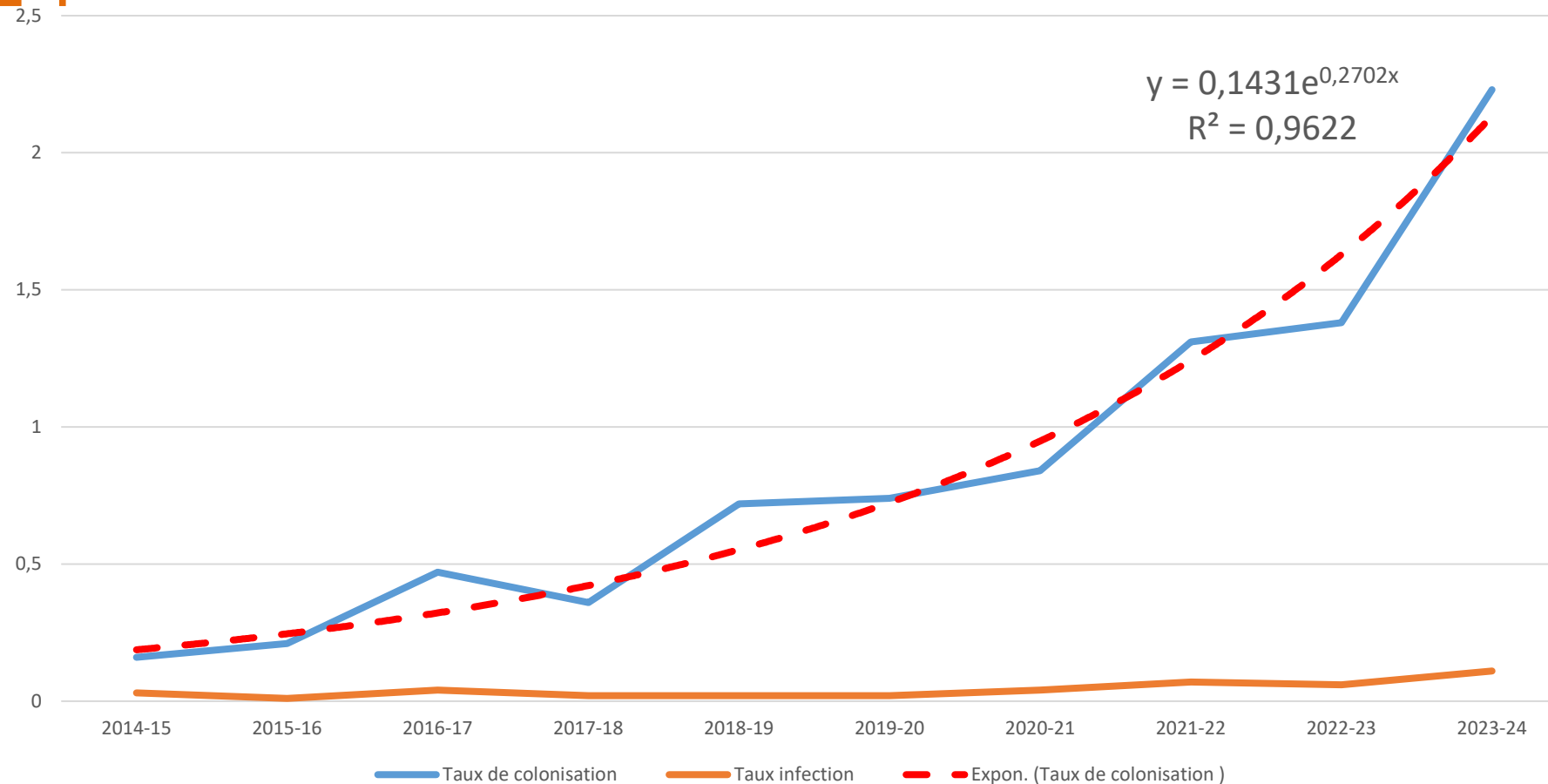
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Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



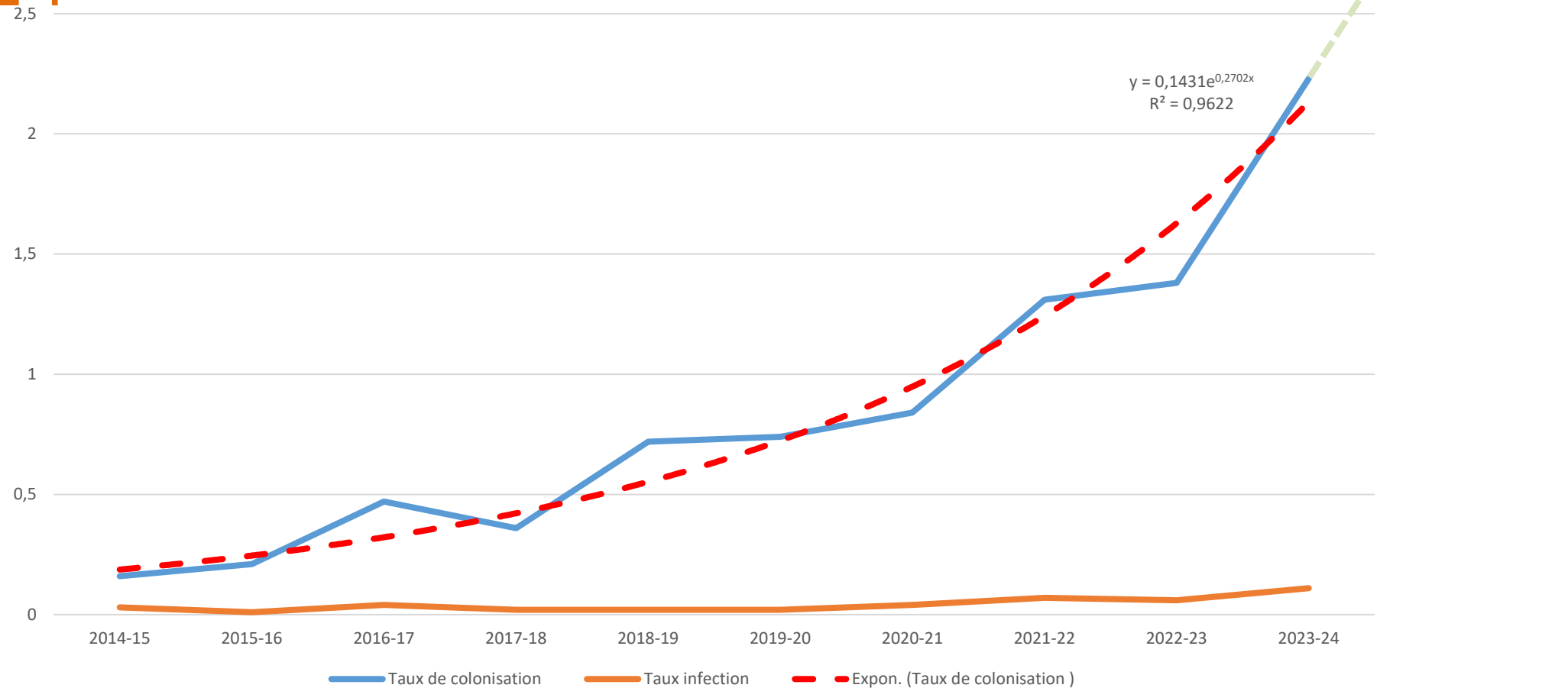
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Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



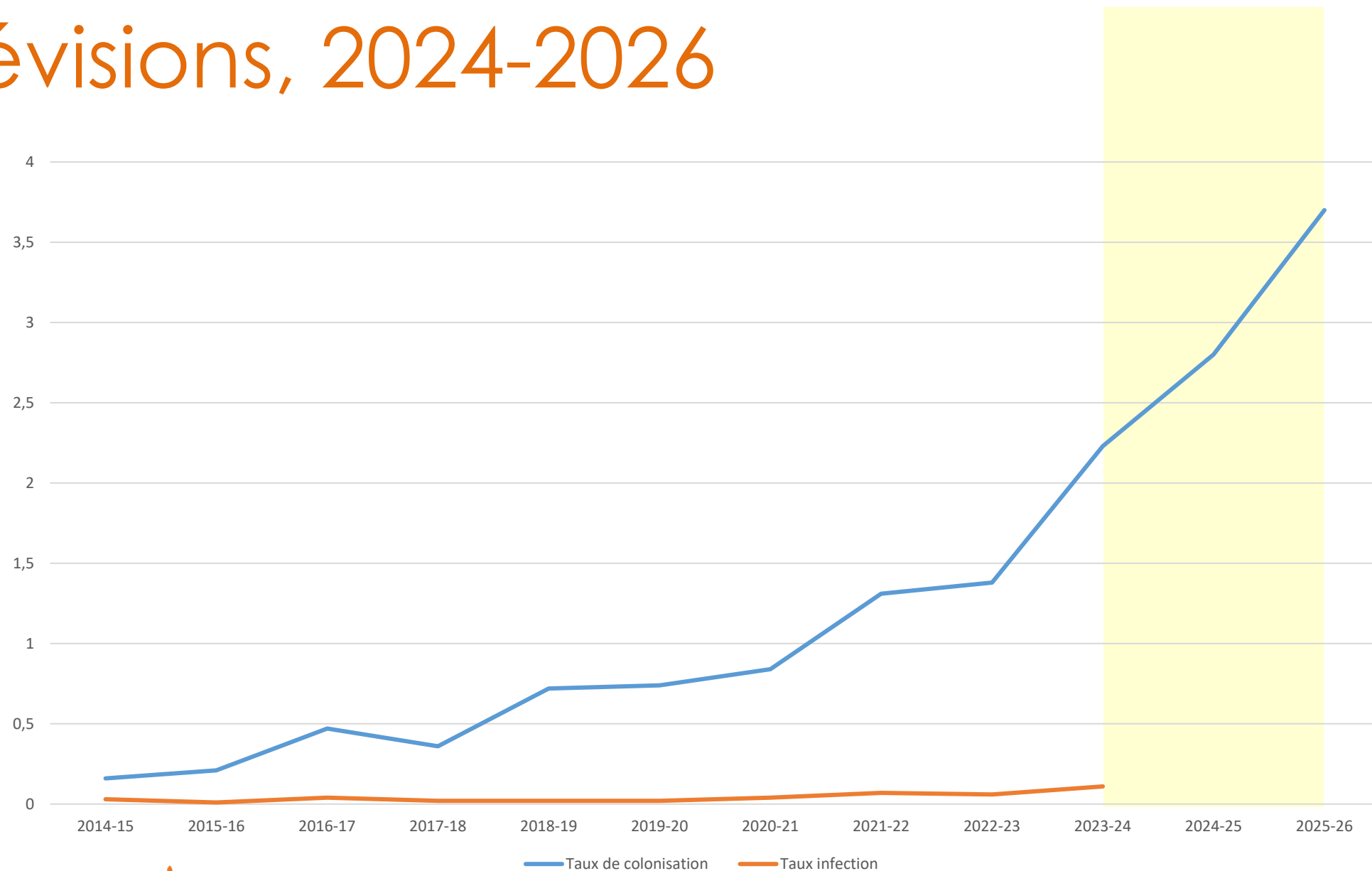
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Evolution du taux d'infection et de colonisation au Québec, 2014-2015 à 2023-2024



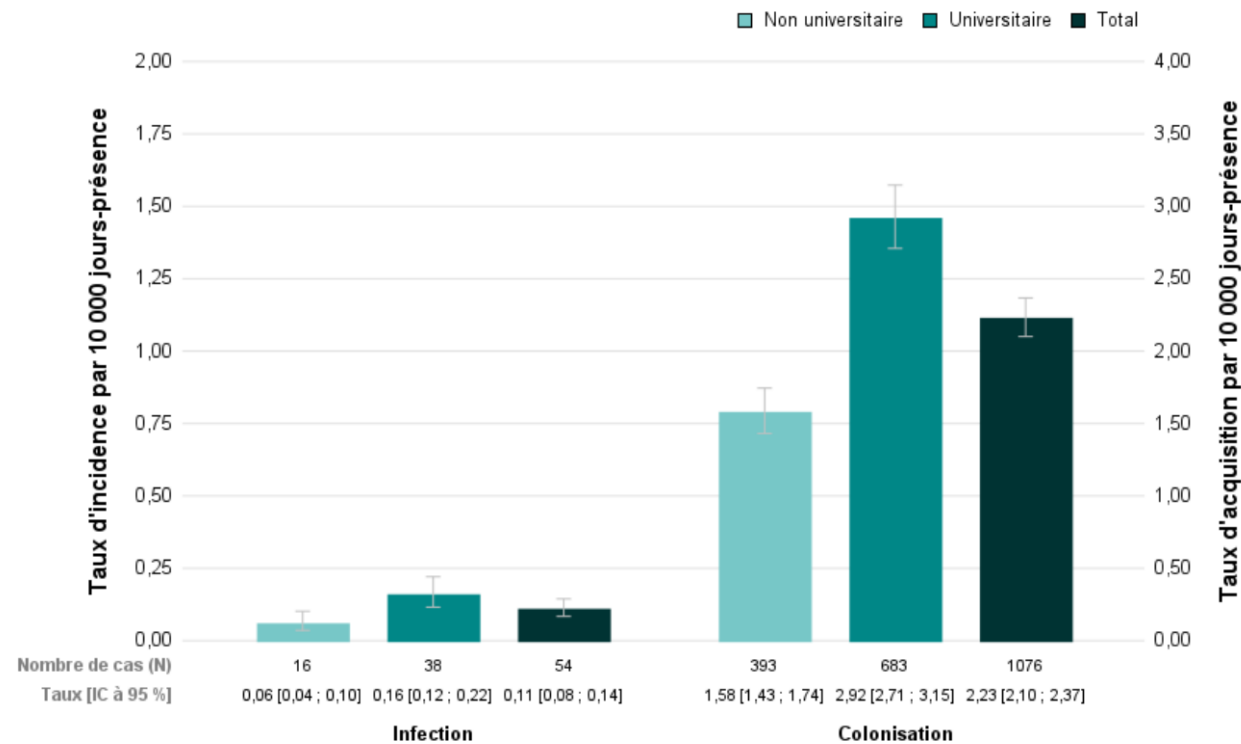
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Prévisions, 2024-2026



Impact de la mission universitaire

Figure 1 Taux d'incidence des infections nosocomiales à BGNPC (cat. 1a et 1b) et taux d'acquisition des colonisations (cat. 1a et 1b) pour les installations participantes (N = 88), 2023-2024



<https://www.inspq.qc.ca/sites/default/files/publications/3577-infections-bacilles-gram-negatif-carbapenemases-QC-2023-2024.pdf>

Figure 2 Évolution du nombre d'installations participantes selon le nombre de colonisations nosocomiales à BGNPC (cat. 1 a et 1 b) déclarés, 2019-2020 à 2023-2024

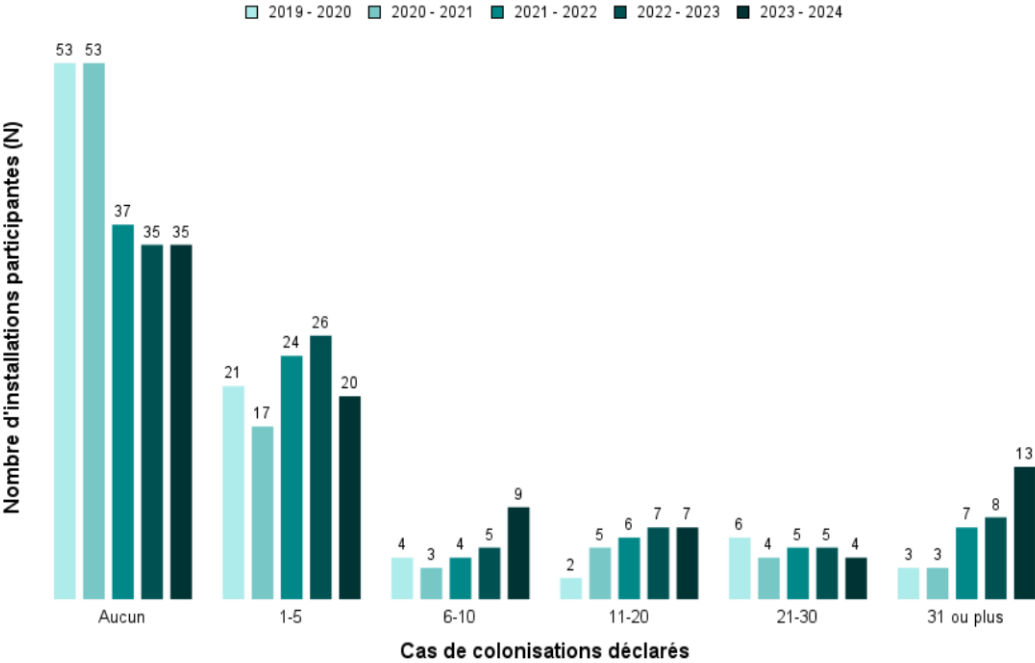
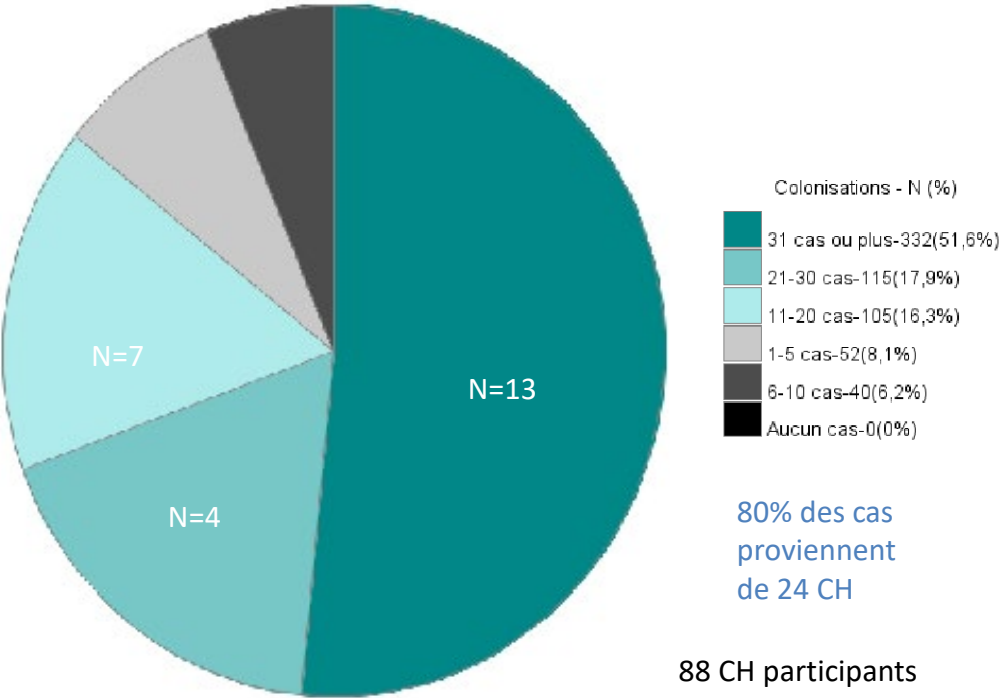
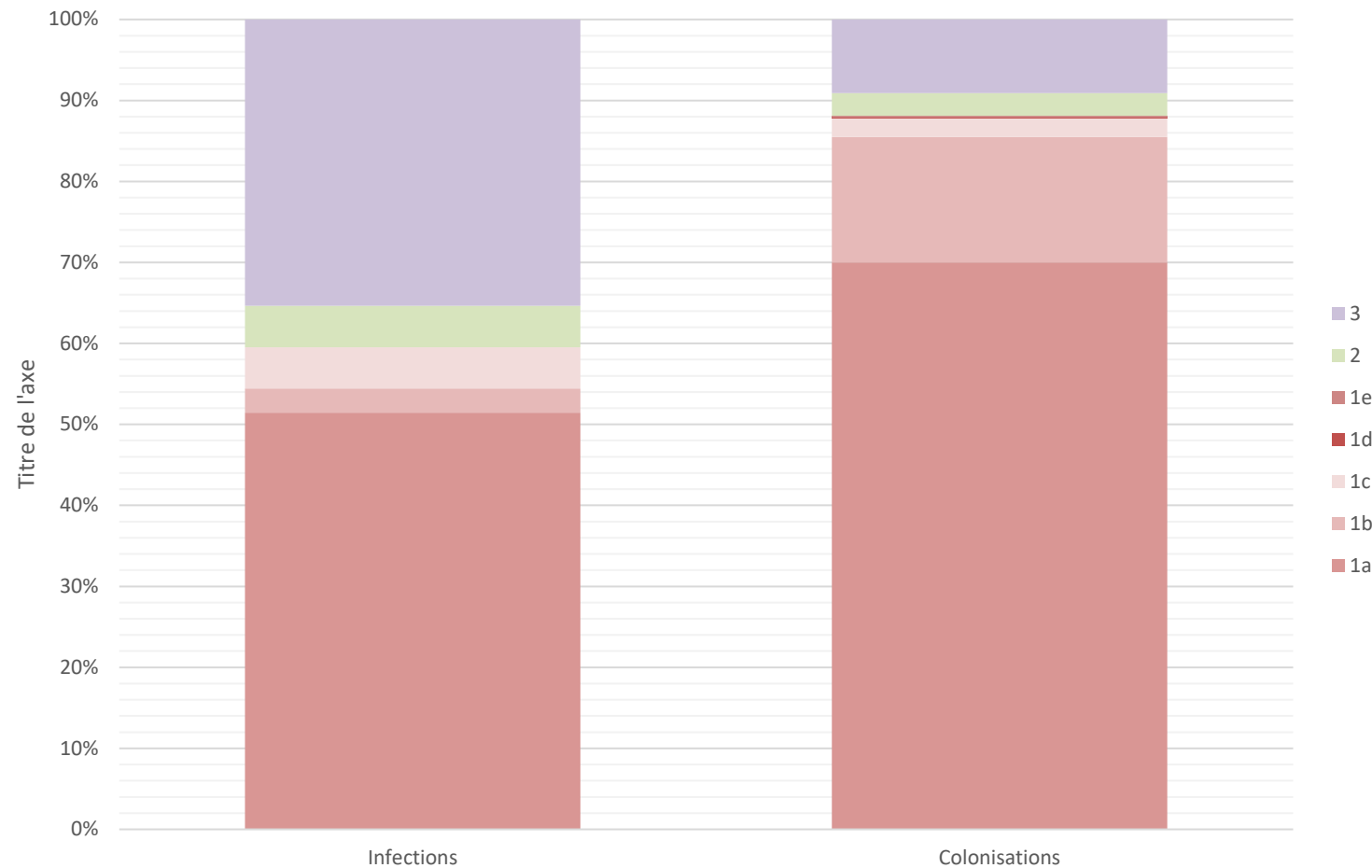


Figure 3 Répartition du nombre de colonisations nosocomiales à BGNPC (cat. 1 a et 1 b) pour les installations participantes (N = 88), 2023-2024



Origine des cas



<https://www.inspq.qc.ca/sites/default/files/publications/3577-infections-bacilles-gram-negatif-carbapenemases-QC-2023-2024.pdf>

INFECTIONS

Tableau 3 Nombre de cas d’infection, selon le type d’infection, et nombre de bactériémies à BGNPC de toute catégorie d’attribution (N = 99), 2023-2024

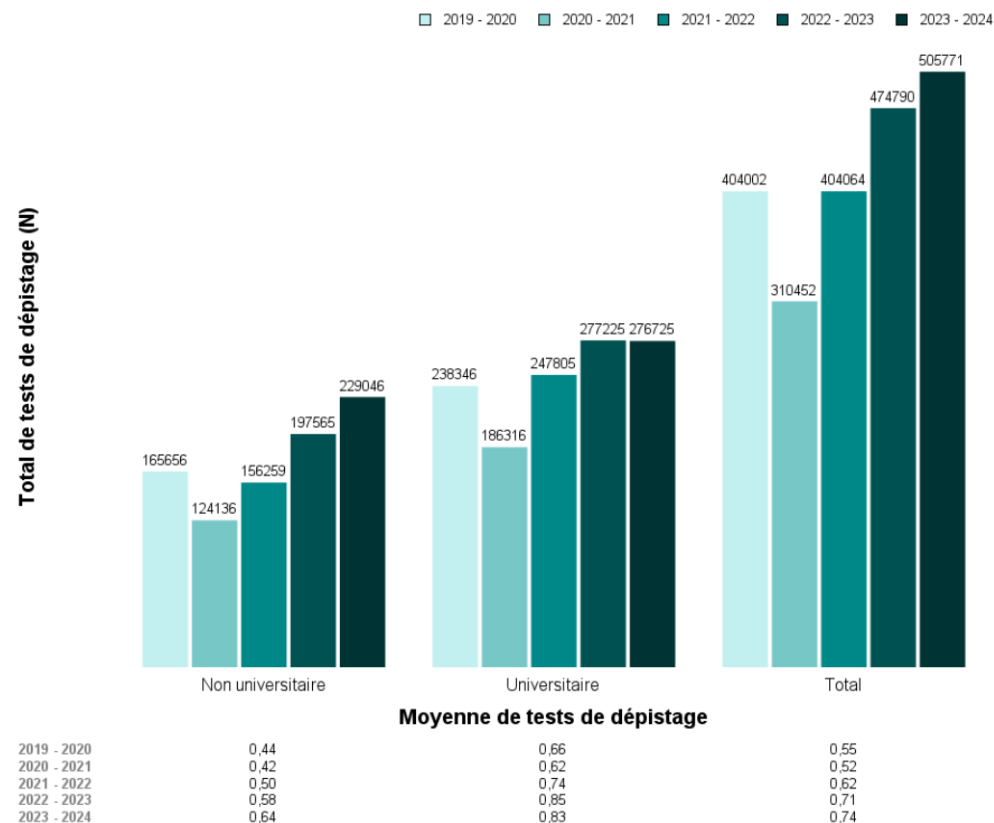
Catégories d’infection	Types d’infection	Infections (N)	Bactériémies secondaires (N)
Bactériémies primaires	BBM	0	-
	BAC	4	-
	Non-BAC	4	-
	HD	0	-
Infections primaires	Urinaire	48	7
	Abdominale	7	2
	Pulmonaire	16	3
	Infection de site opératoire	5	2
	Peau et tissus mous	8	0
	Os et articulations	6	1
	Autres	1	1
Total		99	16

24 bactériémies

Un total de 13 décès de toutes causes (associés ou non aux infections) est observé dans les 30 jours suivant le début de l’infection en 2023-2024, pour une létalité de 13,1 % (tableau 4). En plus des données présentées ci-dessous, onze cas de décès sur les treize sont de catégories (1a, 1b et 1c).

Tests de dépistage

Figure 4 Évolution du nombre total et de la moyenne de tests de dépistage par admission, 2019-2020 à 2023-2024





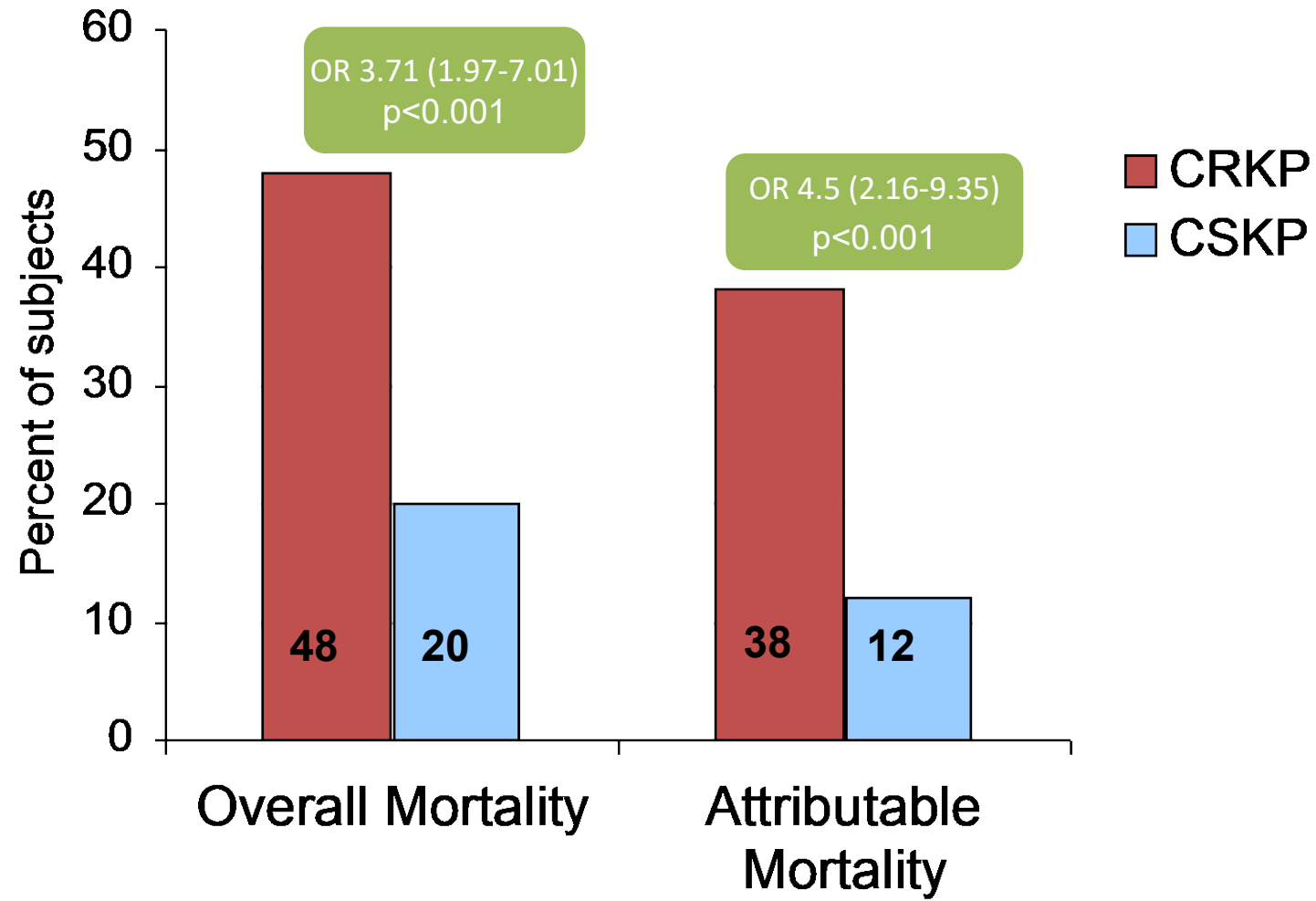
IMPACT CLINIQUE

POURQUOI S'INQUIÉTER ?

- EPC sont dangereux
 - Plus difficile à traiter
 - Peut être plus virulent
 - Mortalité augmentée
 - Morbidité augmentée
- Gourmand en ressources
 - Antibiotiques plus coûteux
 - Durée de l'hospitalisation
 - Mesures d'isolement
- Problèmes dérivés
 - Toxicité des médicaments
 - Qualité des soins en raison de l'isolement dans une chambre individuelle



Mortalité



BMJ Open Impact of carbapenem resistance on mortality in patients infected with *Enterobacteriaceae*: a systematic review and meta-analysis

Ruyin Zhou ¹, Xiangming Fang,^{1,2} Jinjin Zhang,¹ Xiaodong Zheng,³ Shuangyue Shangguan,¹ Shibo Chen,⁴ Yingbo Shen,⁵ Zhihai Liu,⁶ Juan Li,⁷ Rong Zhang,⁸ Jianzhong Shen,⁹ Timothy R Walsh,¹⁰ Yang Wang⁹

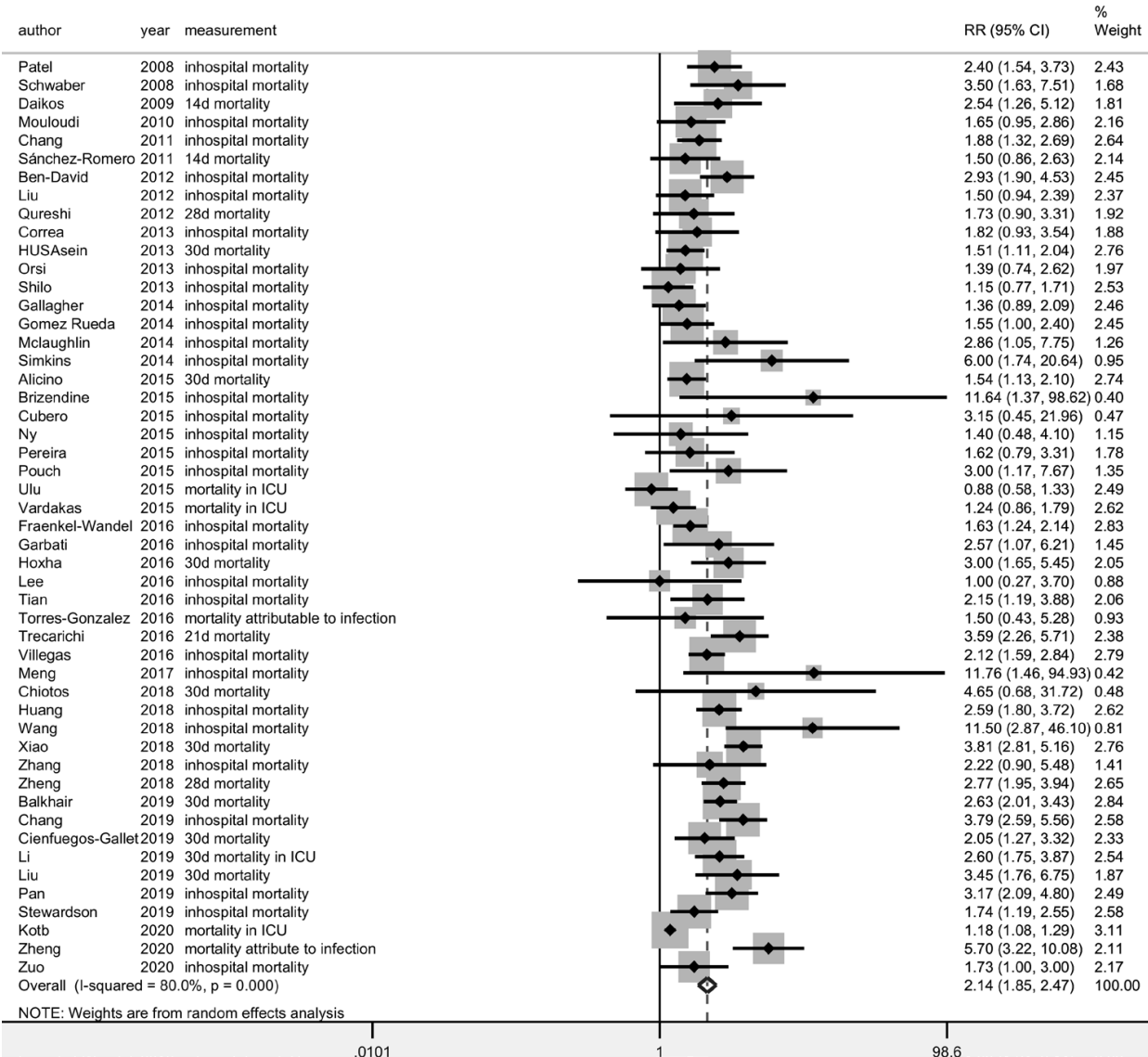


Figure 2 Forest plot of overall mortality in patients with carbapenem-resistant *Enterobacteriaceae* (CRE) versus carbapenem-susceptible *Enterobacteriaceae* (CSE) infections (outcome measure=relative risk). ICU, intensive care unit.

BMJ Open Impact of carbapenem resistance on mortality in patients infected with *Enterobacteriaceae*: a systematic review and meta-analysis

Ruyin Zhou ¹, Xiangming Fang,^{1,2} Jinjin Zhang,¹ Xiaodong Zheng,³ Shuangyue Shangguan,¹ Shibo Chen,⁴ Yingbo Shen,⁵ Zhihai Liu,⁶ Juan Li,⁷ Rong Zhang,⁸ Jianzhong Shen,⁹ Timothy R Walsh,¹⁰ Yang Wang⁹

Table 2 Pool estimated results for different types of mortality outcome

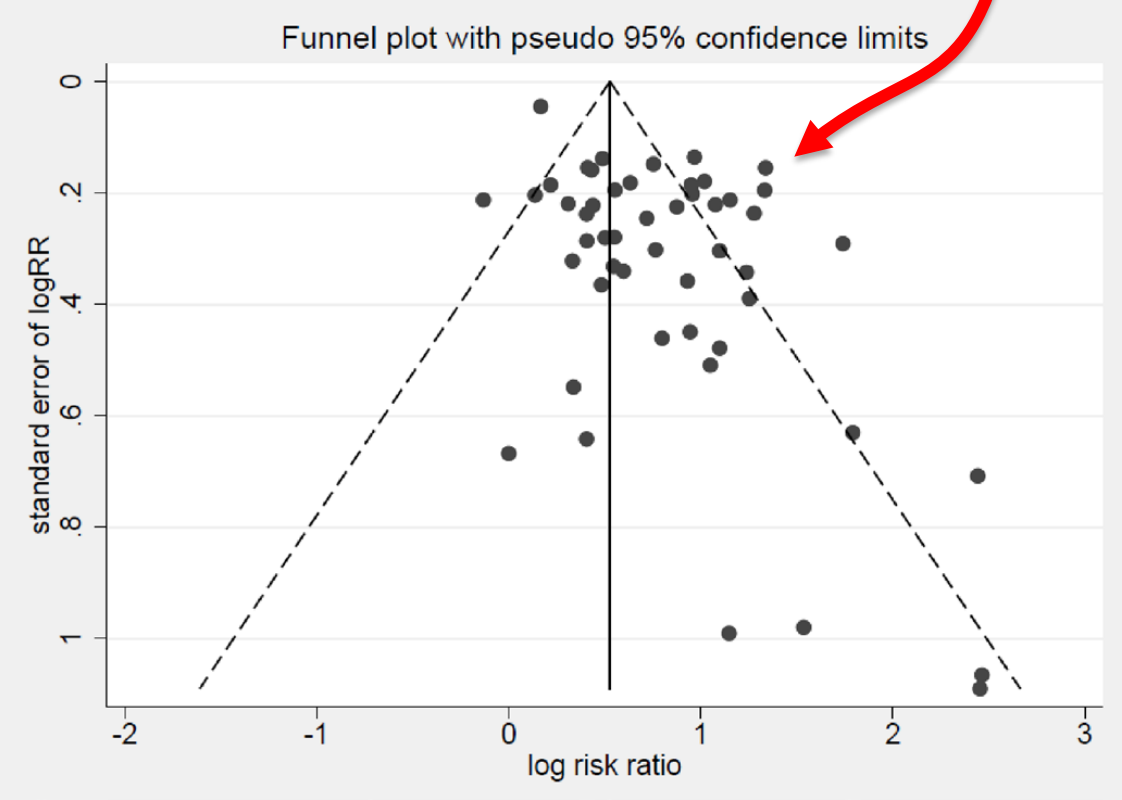
	Number	Number	Unweighted means of	Unweighted means of	P value
Mortality outcome	Number of ORE	of OSE	mortality among ORE	mortality among OSE	(significance)
type	studies	patients	patients	patients	(test of H0=1)
In-hospital mortality	31	1098	37.93	42.30%	20.00%
				2.09 (1.91 to 2.27)	0.000
28-day or 30-day mortality	17	1101	24.63	42.85%	19.88%
				2.23 (1.83 to 2.71)	0.000
21-day mortality	1	161	117	52.20%	14.50%
				3.39 (2.29 to 5.71)	0.000
14-day mortality	4	84	287	45.89%	27.81%
				1.70 (1.24 to 2.35)	0.001
6-day or 7-day mortality	3	149	372	25.80%	6.57%
				3.89 (2.29 to 6.63)	0.000
Mortality attributable to infection	8	391	778	43.30%	17.45%
				2.71 (1.97 to 3.61)	0.000
Mortality in ICU	4	1065	824	59.83%	50.50%
				1.17 (1.00 to 1.29)	0.000
30-day mortality in ICU	1	244	263	29.80%	11.00%
				2.50 (1.50 to 3.87)	0.000

ORE, carbapenem-resistant *Enterobacteriaceae*; OSE, carbapenem-susceptible *Enterobacteriaceae*; ICU, intensive care unit.

BMJ Open Impact of carbapenem resistance on mortality in patients infected with *Enterobacteriaceae*: a systematic review and meta-analysis

Ruyin Zhou ¹, Xiangming Fang,^{1,2} Jinjin Zhang,¹ Xiaodong Zheng,³ Shuangyue Shangguan,¹ Shibo Chen,⁴ Yingbo Shen,⁵ Zhihai Liu,⁶ Juan Li,⁷ Rong Zhang,⁸ Jianzhong Shen,⁹ Timothy R Walsh,¹⁰ Yang Wang⁹

CRP carbapenem-resistant <i>Enterobacteriaceae</i> ; CSF carbapenem-susceptible <i>Enterobacteriaceae</i> ; ICU, intensive care unit.															
CRP carbapenem-resistant <i>Enterobacteriaceae</i>		CSF carbapenem-susceptible <i>Enterobacteriaceae</i>		ICU, intensive care unit											
14-day mortality	4	84	287	45.00%	27.00%	1.70 (1.24 to 2.35)	0.001	6-day or 7-day mortality	3	149	372	25.00%	6.57%	3.60 (2.20 to 5.83)	0.000
Mortality attributable to infection	8	391	778	43.00%	17.45%	2.74 (1.97 to 3.81)	0.000	30-day mortality in ICU	4	1065	824	59.83%	50.50%	1.17 (1.00 to 1.28)	0.000
30-day mortality in ICU	1	244	263	29.00%	11.00%	2.80 (1.50 to 5.27)	0.000								



Léger biais de publication
(manque de petites études
qui ne détectent pas de
mortalité augmentée)

Figure 4 Funnel plot of studies evaluating mortality of patients with infections due to carbapenem-resistant compared with carbapenem-susceptible *Enterobacteriaceae*. Zhou R et al. *BMJ Open* 2021;11:e054971.

“A post-antibiotic era means, in effect, an end to modern medicine as we know it. Things as common as strep throat or a child’s scratched knee could once again kill.”

**-Dr. Margaret Chan, Director General of the World Health Organization
Keynote Address, Conference On Combating Antimicrobial Resistance,
Copenhagen, Denmark (March 14, 2012)**



BAD BUGS, NO DRUGS

As Antibiotic Discovery Stagnates ...
A Public Health Crisis Brews

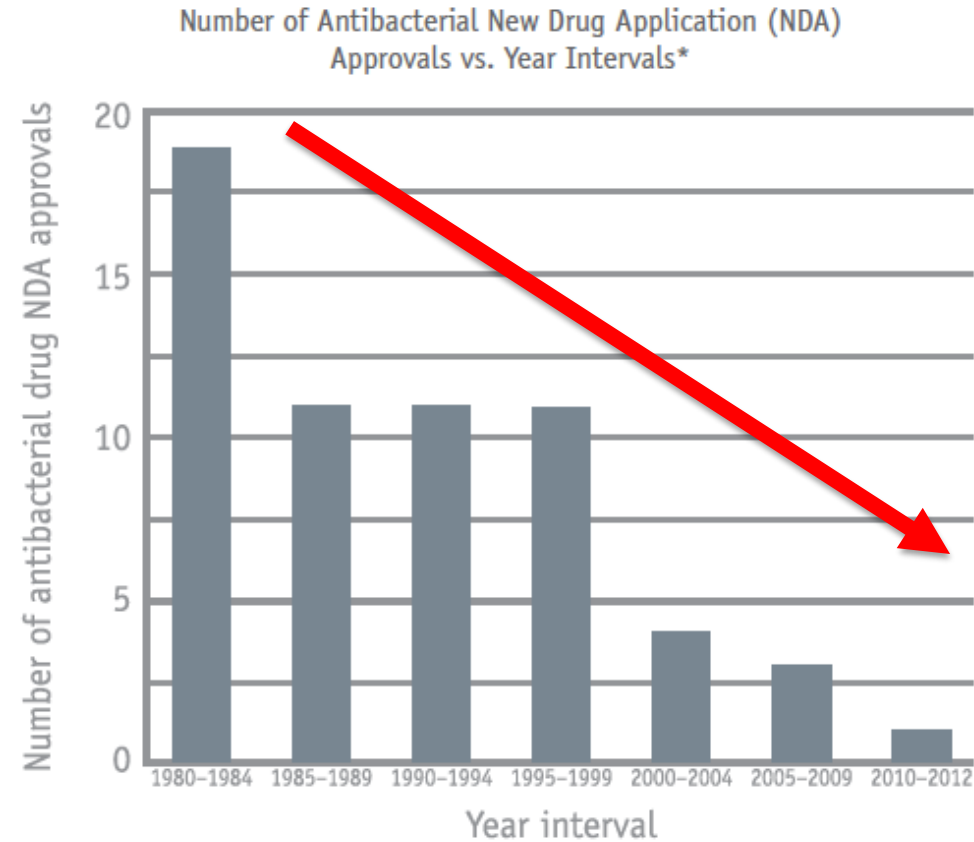


 **IDSA**
Infectious Diseases Society of America

July 2004

Tomorrow's Antibiotics: The Drug Pipeline

The number of new antibiotics developed and approved has steadily decreased in the past three decades, leaving fewer options to treat resistant bacteria.



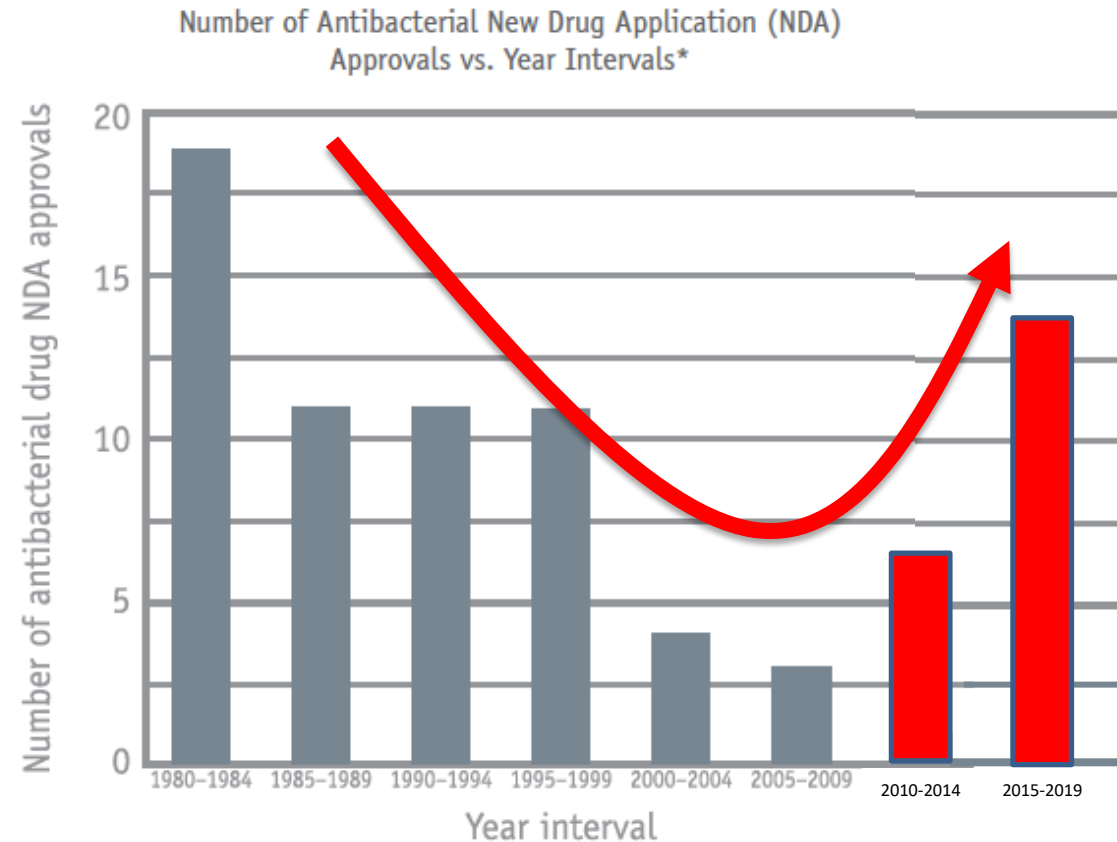
*Intervals from 1980-2009 are 5-year intervals; 2010-2012 is a 3-year interval. Drugs are limited to systemic agents. Data courtesy of FDA's Center for Drug Evaluation and Research (CDER).

<http://www.cdc.gov/drugresistance/threat-report-2013/>

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Cefiderocol 2019
Imipenem rele: 2019
Lefalumin: 2019
Seracycline: 2018
Plazomicin: 2018
Eravacycline: 2018
Ryfamicine: 2018
Omdacycline: 2018
Meropenem-vabor: 2017
Secnidazole: 2017
Delafloxacin: 2017
Ozenoxacin: 2017
Ceftazidime-avib: 2015
Dalbavancin: 2014
Ceftolozane/tazo: 2014
Finafloxacin: 2014
Tedilozid: 2014
Ceftaroline: 2010
Fidaxomycin 2011

<http://www.cdc.gov/drugresistance/threat-report-2013/>

<https://pubs.acs.org/doi/10.1021/acs.jmedchem.0c01516>

Infectious Diseases Society of America 2024 Guidance on the Treatment of Antimicrobial-Resistant Gram-Negative Infections

Rouge : disponible uniquement par l'entremise de SAP au Canada

Pranita D. Tamma,^{1,6} Emily L. Heil,² Julie Ann Justo,³ Amy J. Mathers,⁴ Michael J. Satlin,⁵ and Robert A. Bonomo⁶

¹Department of Pediatrics, Johns Hopkins University School of Medicine, Baltimore, Maryland, USA; ²Department of Practice, Sciences, and Health-Outcomes Research, University of Maryland School of Pharmacy, Baltimore, Maryland, USA; ³Department of Pharmacy, Dartmouth Hitchcock Medical Center, Lebanon, New Hampshire, USA; ⁴Departments of Medicine and Pathology, University of Virginia, Charlottesville, Virginia, USA; ⁵Department of Medicine, Weill Cornell Medicine, New York, New York, USA; and ⁶Medical Service and Center for Antimicrobial Resistance and Epidemiology, Louis Stokes Cleveland Veterans Affairs Medical Center, University Hospitals Cleveland Medical Center and Departments of Medicine, Pharmacology, Molecular Biology, and Microbiology, Case Western Reserve University, Cleveland, Ohio, USA

- Infection due à KPC: Meropenem-vaborbactam, ceftazidime-avibactam, imipenem-relebactam (option: cefiderocol)
- Infections due à NDM: ceftazidime-avibactam+aztreonam, cefiderocol
- Infections due à Oxa-48: ceftazidime-avibactam, cefiderocol
- Tigecycline: alternative pour infection n'impliquant pas la circulation sanguine ou les voies urinaires
- Colistin : n'est plus suggéré à l'exception de la cystite EPC non compliquée

Bucking the trend... Except in Canada?

Clinical Infectious Diseases

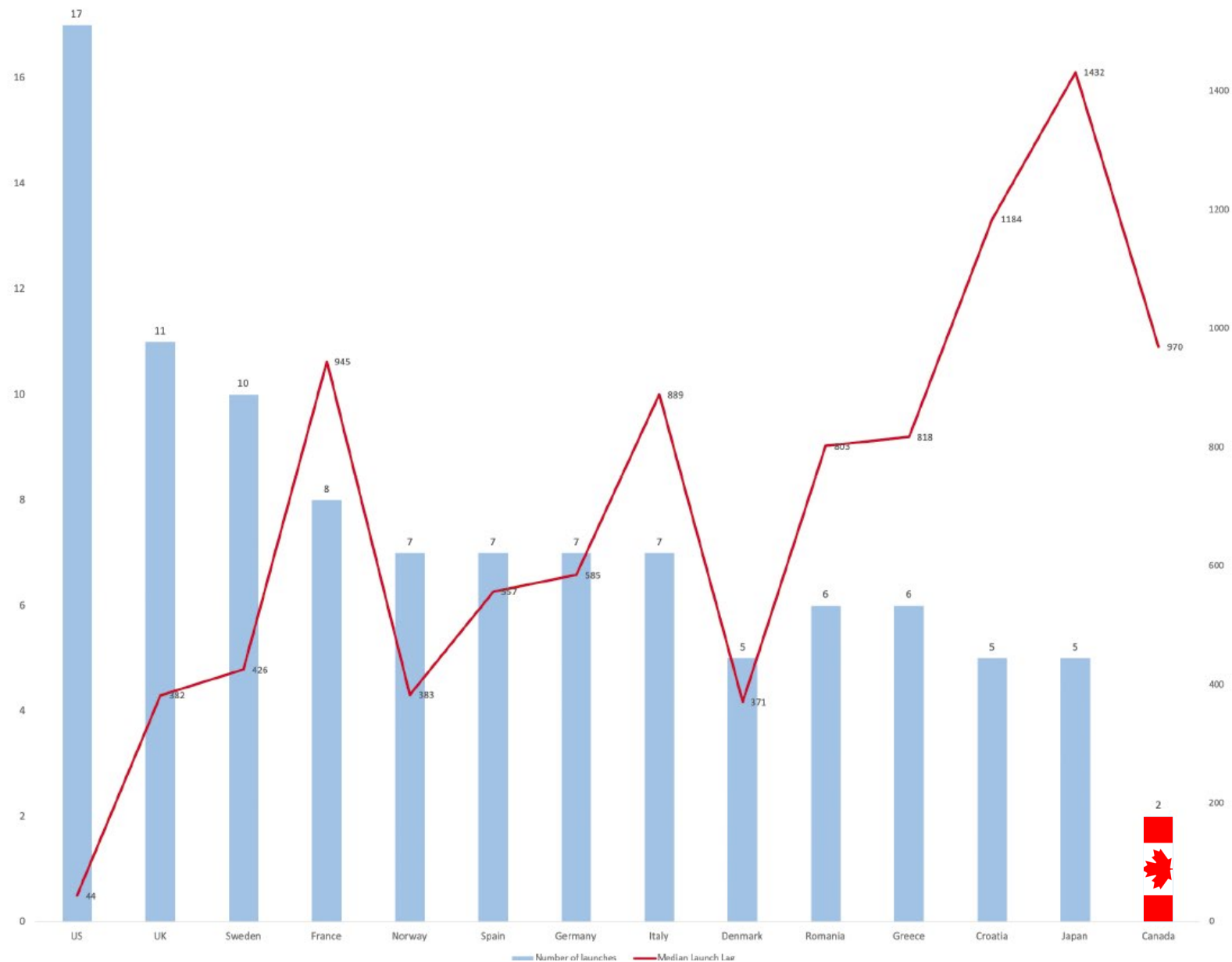
MAJOR ARTICLE



Patient Access in 14 High-Income Countries to New Antibacterials Approved by the US Food and Drug Administration, European Medicines Agency, Japanese Pharmaceuticals and Medical Devices Agency, or Health Canada, 2010–2020

Kevin Outterson,^{1,2} Ebiowei S. F. Orubu,^{3,4} John Rex,^{5,6} Christine Årdal,⁷ and Muhammad H. Zaman⁴

¹Boston University School of Law, Boston, Massachusetts, USA; ²CARB-X, Boston, Massachusetts, USA; ³Social Innovation on Drug Resistance Program, Boston University, Boston, Massachusetts, USA; ⁴Department of Biomedical Engineering, Boston University, Boston, Massachusetts, USA; ⁵F2G Limited, Eccles, Cheshire, United Kingdom; ⁶McGovern Medical School at The University of Texas Health Science Center at Houston, Houston, Texas, USA; and ⁷Antimicrobial Resistance Centre, Norwegian Institute of Public Health, Oslo, Norway



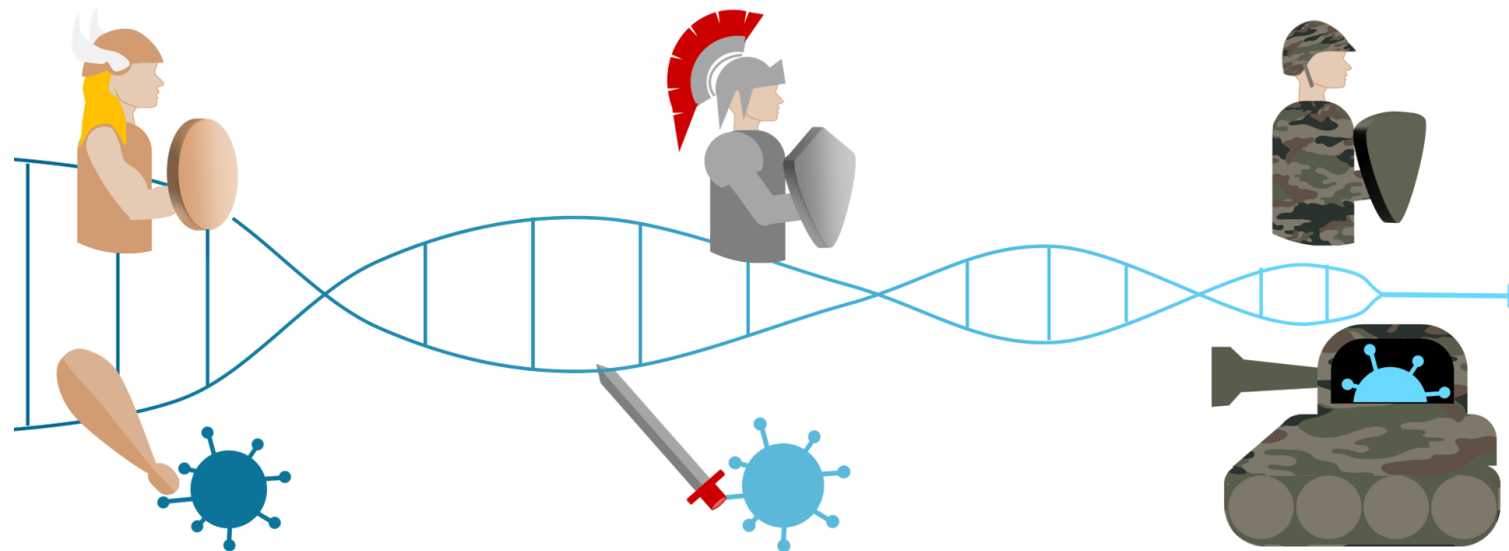
Délais mise en marché: 2.7 years

2/17 (12%) approuvés au CAN

Figure 1. Median launch lag, in days (right axis) and number of launches (left axis) in 14 high-income countries for new molecular entity antibacterials first approved by the United States Food and Drug Administration, European Medicines Agency, Japanese Pharmaceuticals and Medical Devices Agency, or Health Canada, 2010–2019. Abbreviations: UK, United Kingdom; US, United States.

Conclusions

- EPC se propagent de manière **exponentielle**
- Les EPC sont associés à une **morbidity et mortalité** augmentées
- Les **nouveaux traitements** non disponibles au Canada
- **Il est temps d'agir!**



THANK YOU!

