



12 aprile 2025

Meyer Health Campus

Firenze



Proteggere i più piccoli:

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico



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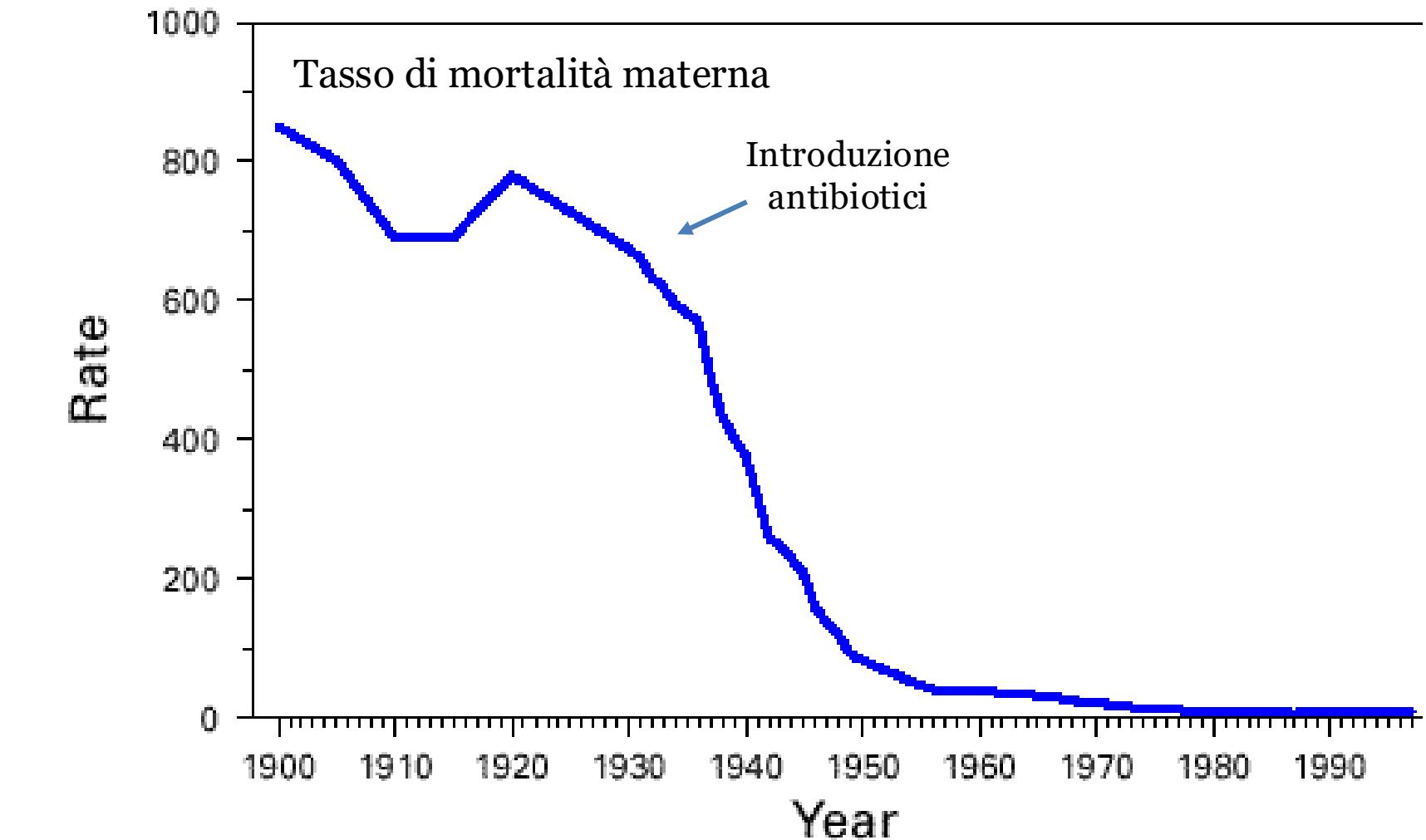
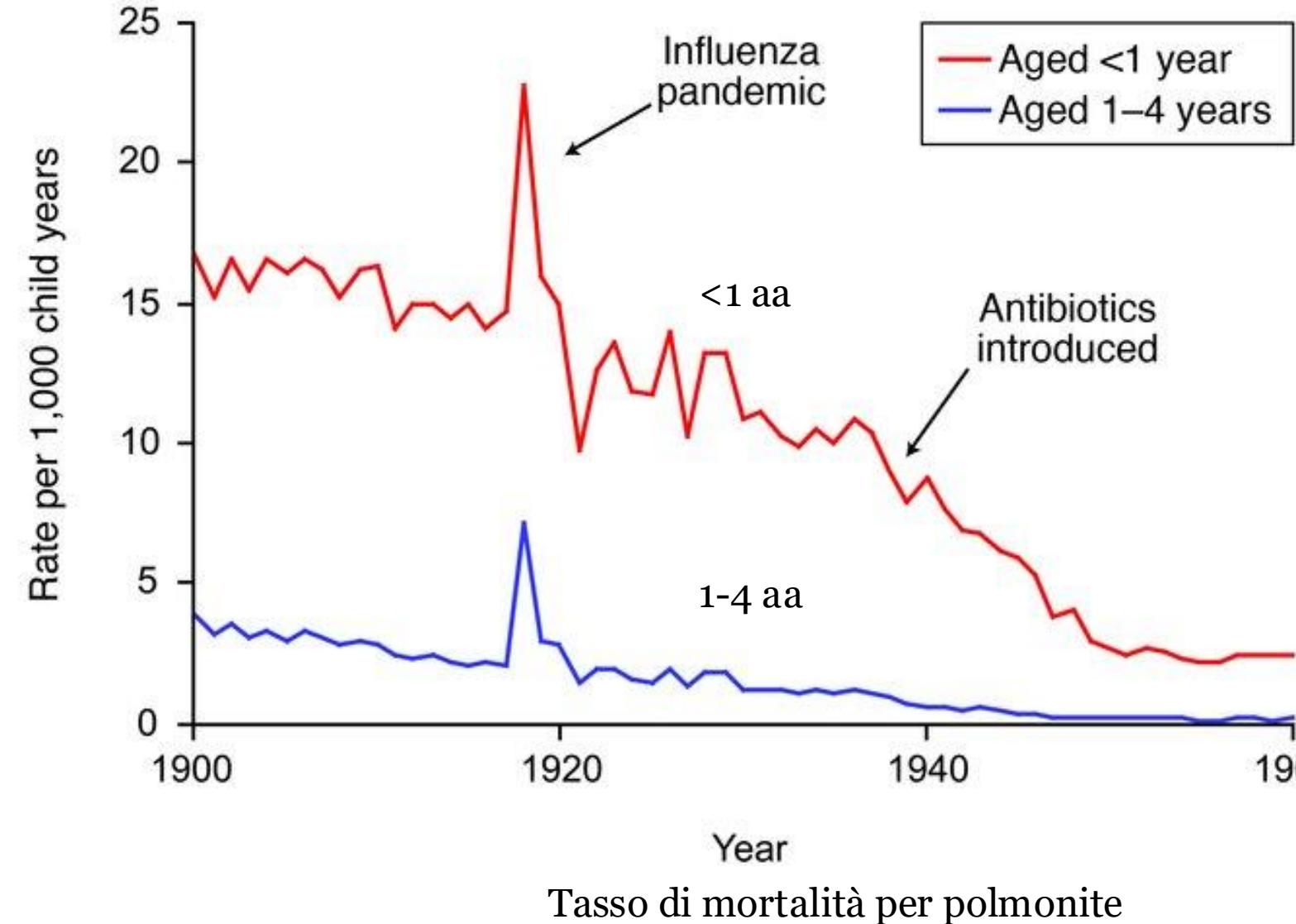
Antimicrobial Stewardship nel bambino

Carlotta Montagnani
SOC Malattie Infettive AOU Meyer IRCCS



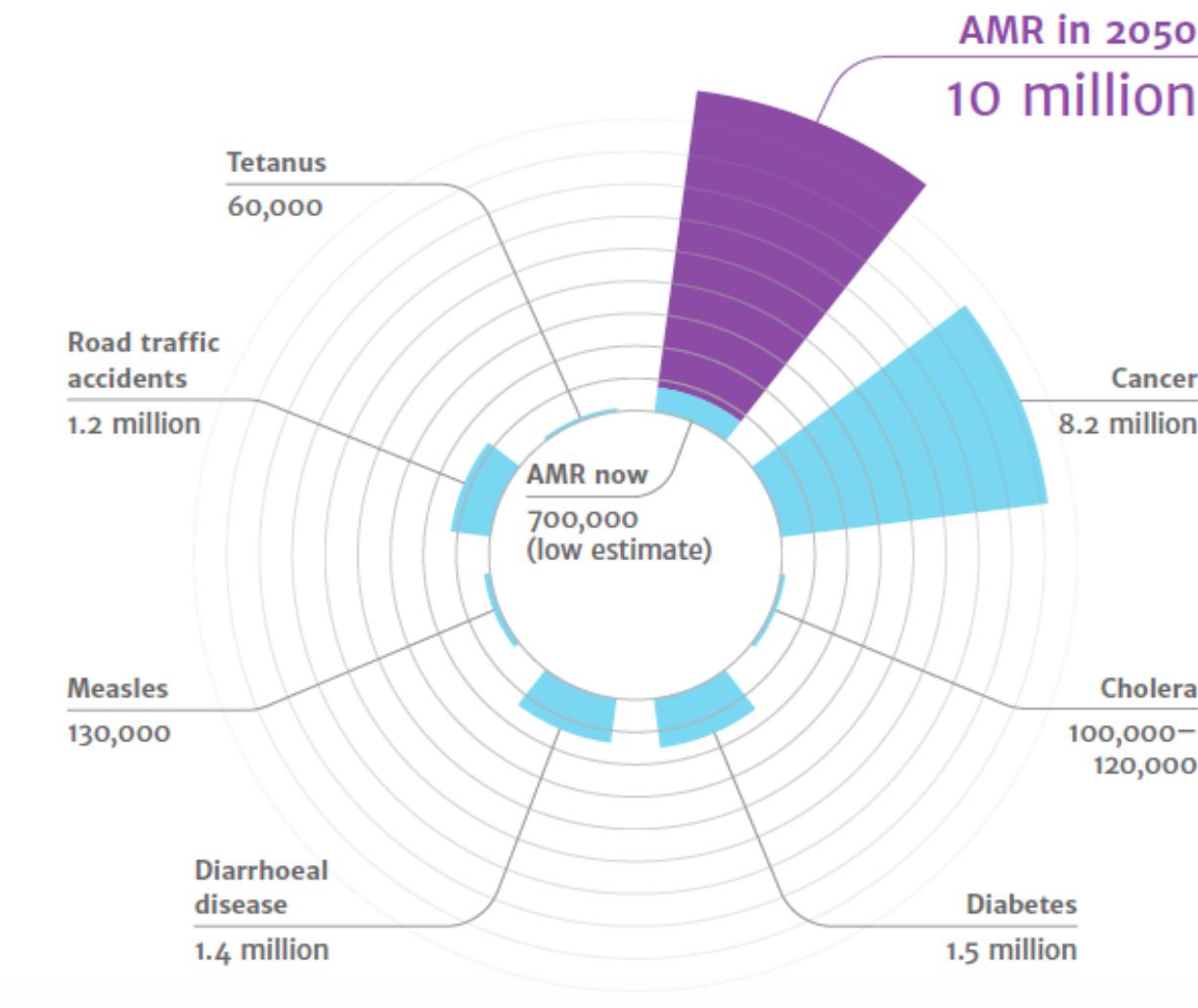
Proteggere i più piccoli:
**Strategie innovative per la
prevenzione delle
infezioni correlate all'assistenza
in ambito pediatrico**

Importanza antibiotici in età pediatrica



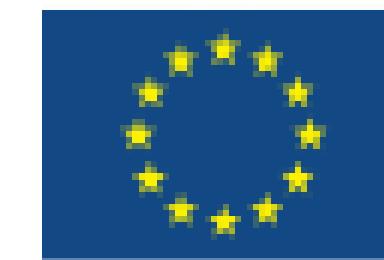
Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

DEATHS ATTRIBUTABLE TO AMR EVERY YEAR



**AMR: a major European
and Global challenge**

Extra healthcare costs and
productivity losses due to
multidrug-resistant bacteria in
the EU:
EUR 1.5 billion each year



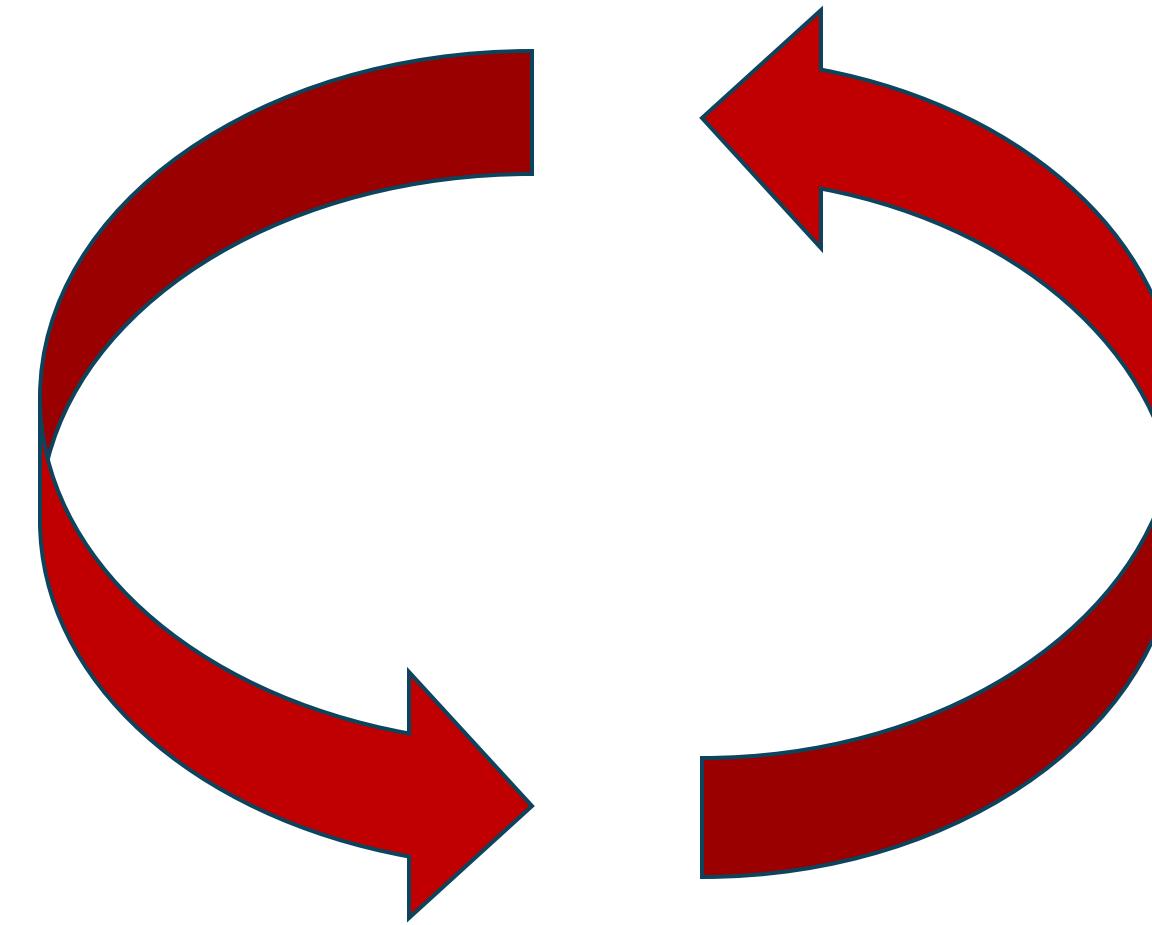
European
Commission



Selection of antibiotic-resistant pathogens in the community

Yagupsky P, Pediatr Infect Dis J 2006;25:974-6

**Selection of
resistant
organisms**



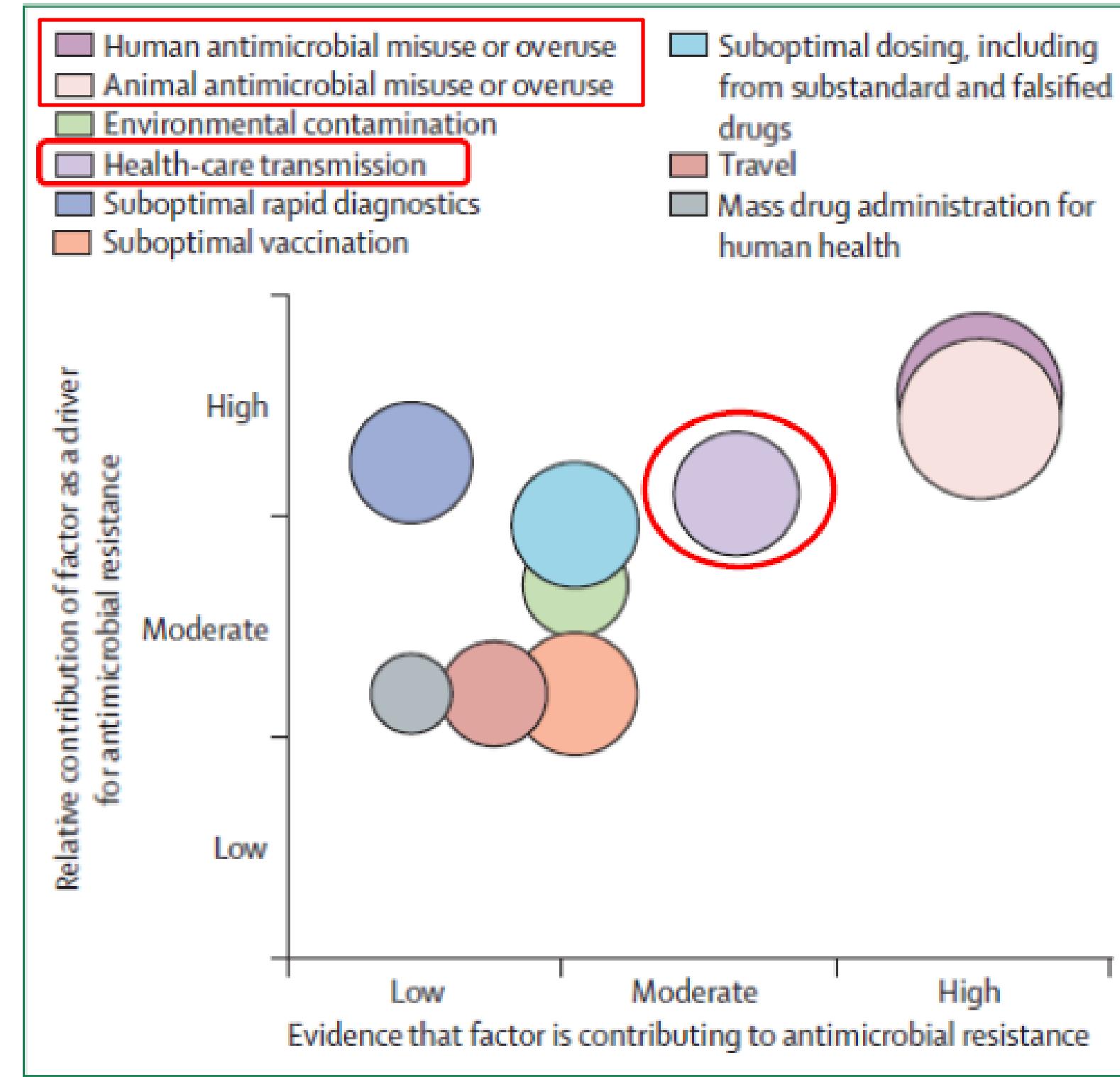
**Antibiotics
overuse**

**Therapeutic
failure**

**Additional
antibiotics
administration**

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Fattori contribuenti l'AMR



Holmes HA, et al. Lancet 2016

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

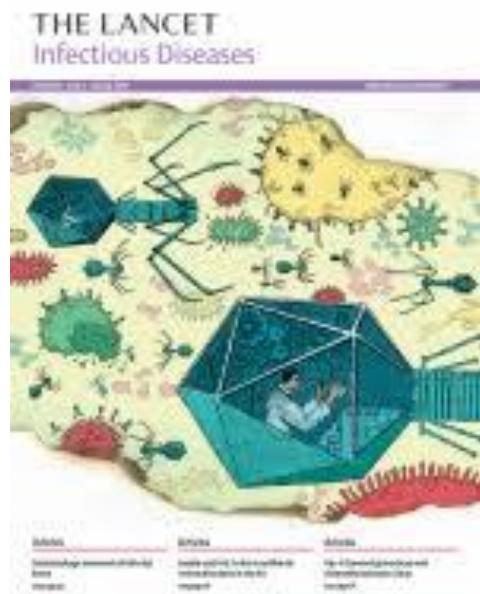
Non solo un problema dell'età adulta...



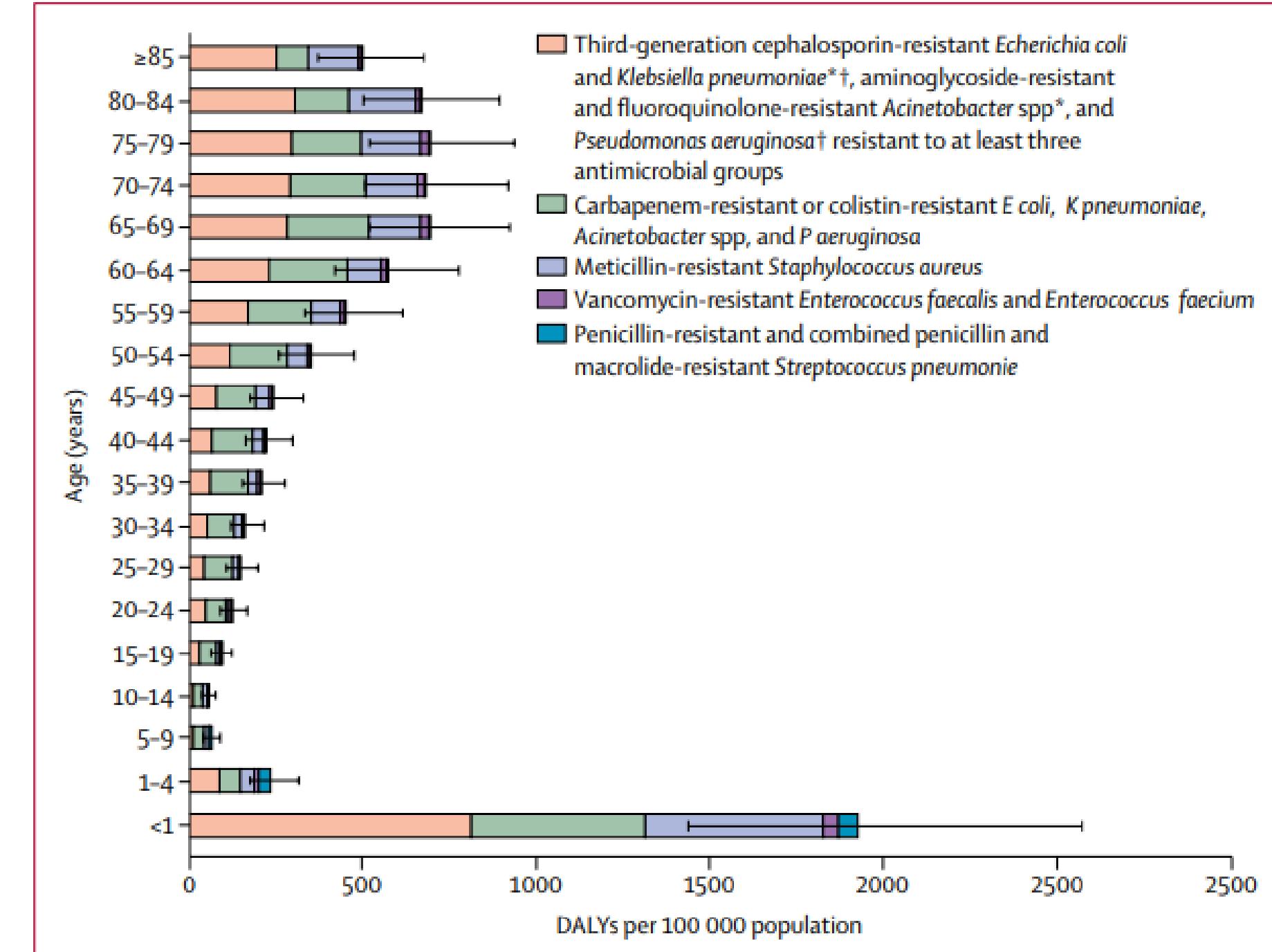
Attributable deaths and disability-adjusted life-years caused by infections with antibiotic-resistant bacteria in the EU and the European Economic Area in 2015: a population-level modelling analysis



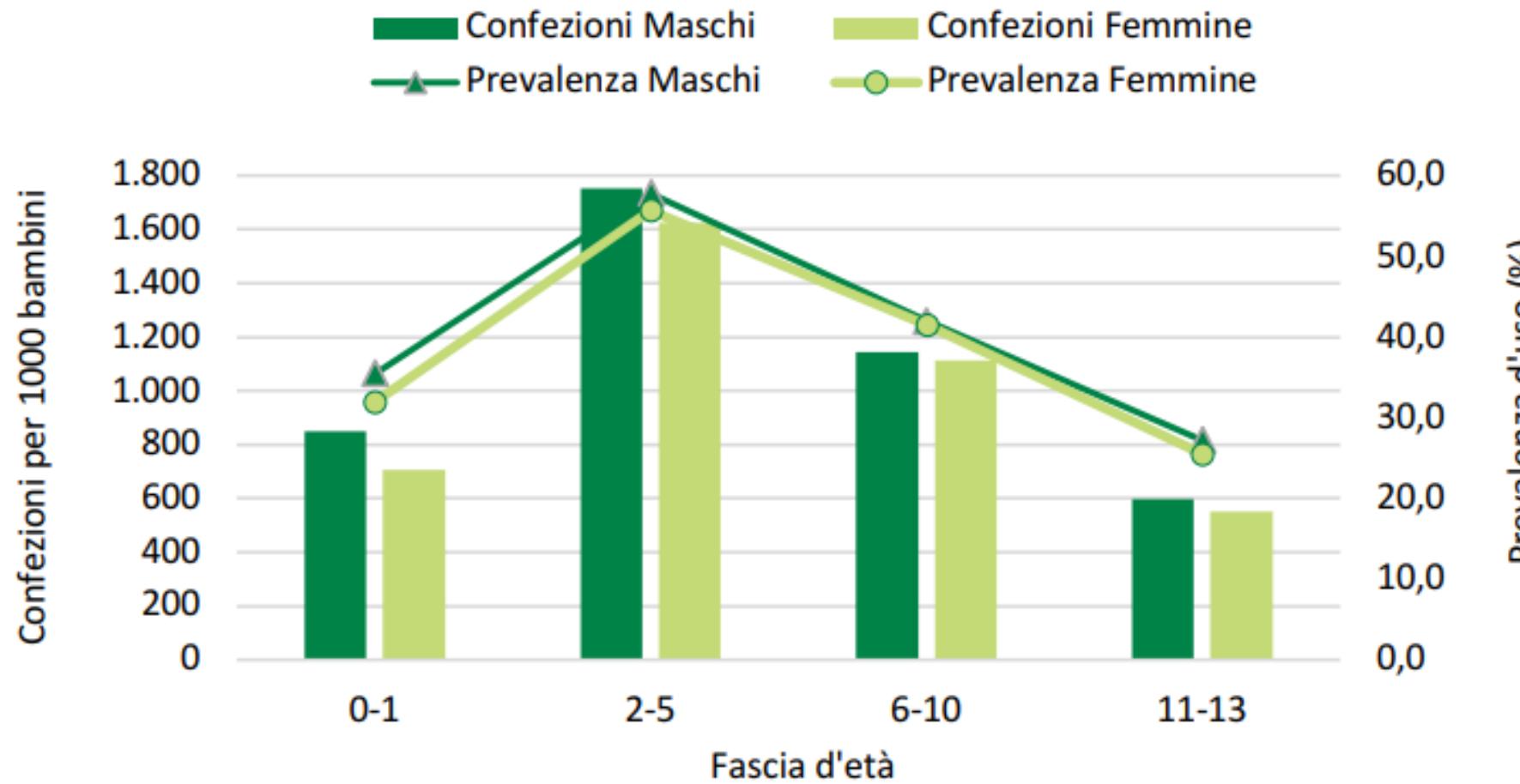
Alessandro Cassini, Liselotte Diaz Höglberg, Diamantis Plachouras, Annalisa Quattrocchi, Ana Hoxha, Gunnar Skov Simonsen, Mélanie Colomb-Cotinat, Mirjam E Kretzschmar, Brecht Devleesschauwer, Michele Cecchini, Driss Ait Ouakrim, Tiago Cravo Oliveira, Marc J Struelens, Carl Suetens, Dominique L Monnet, and the Burden of AMR Collaborative Group*



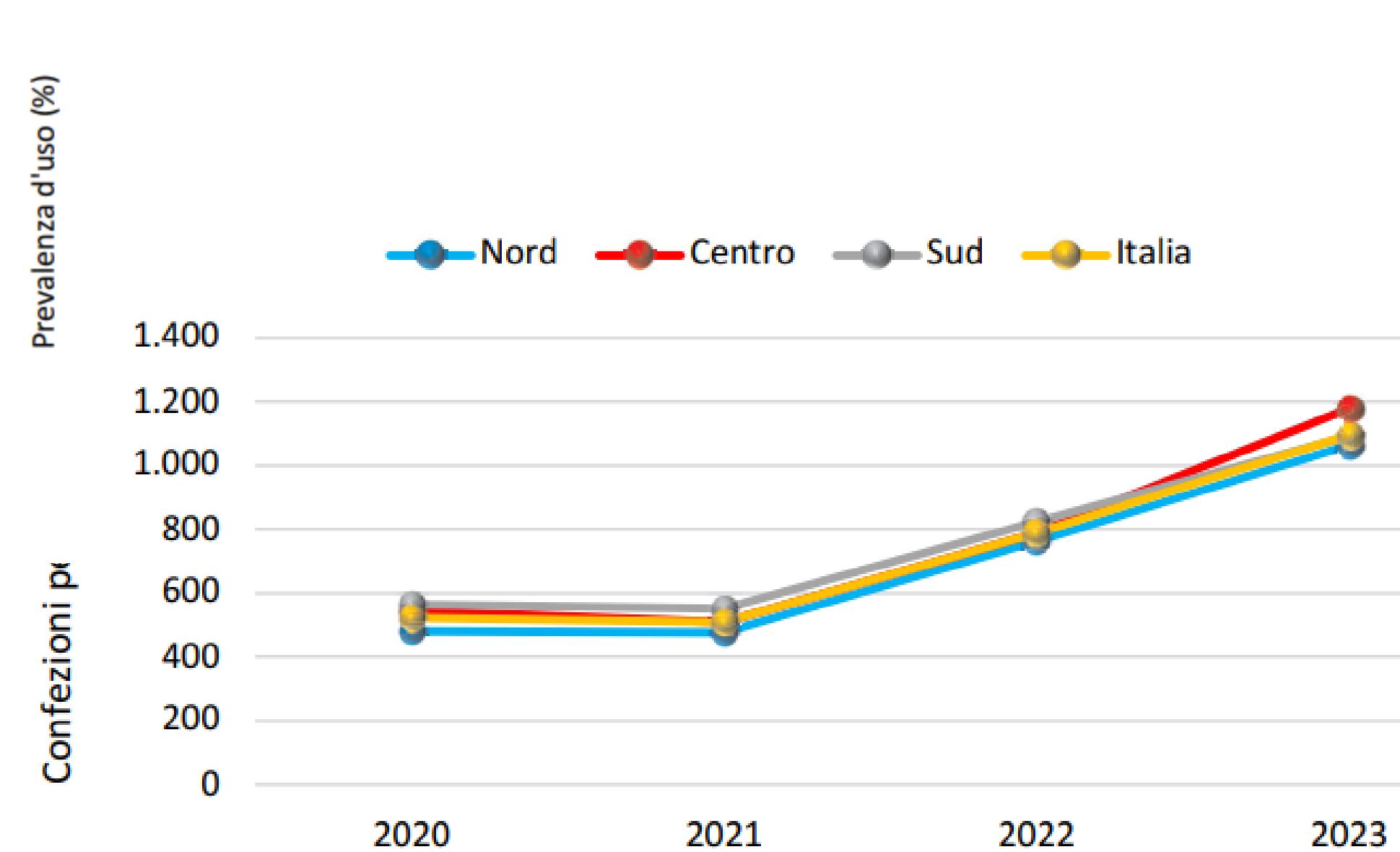
Cassini A, 2019



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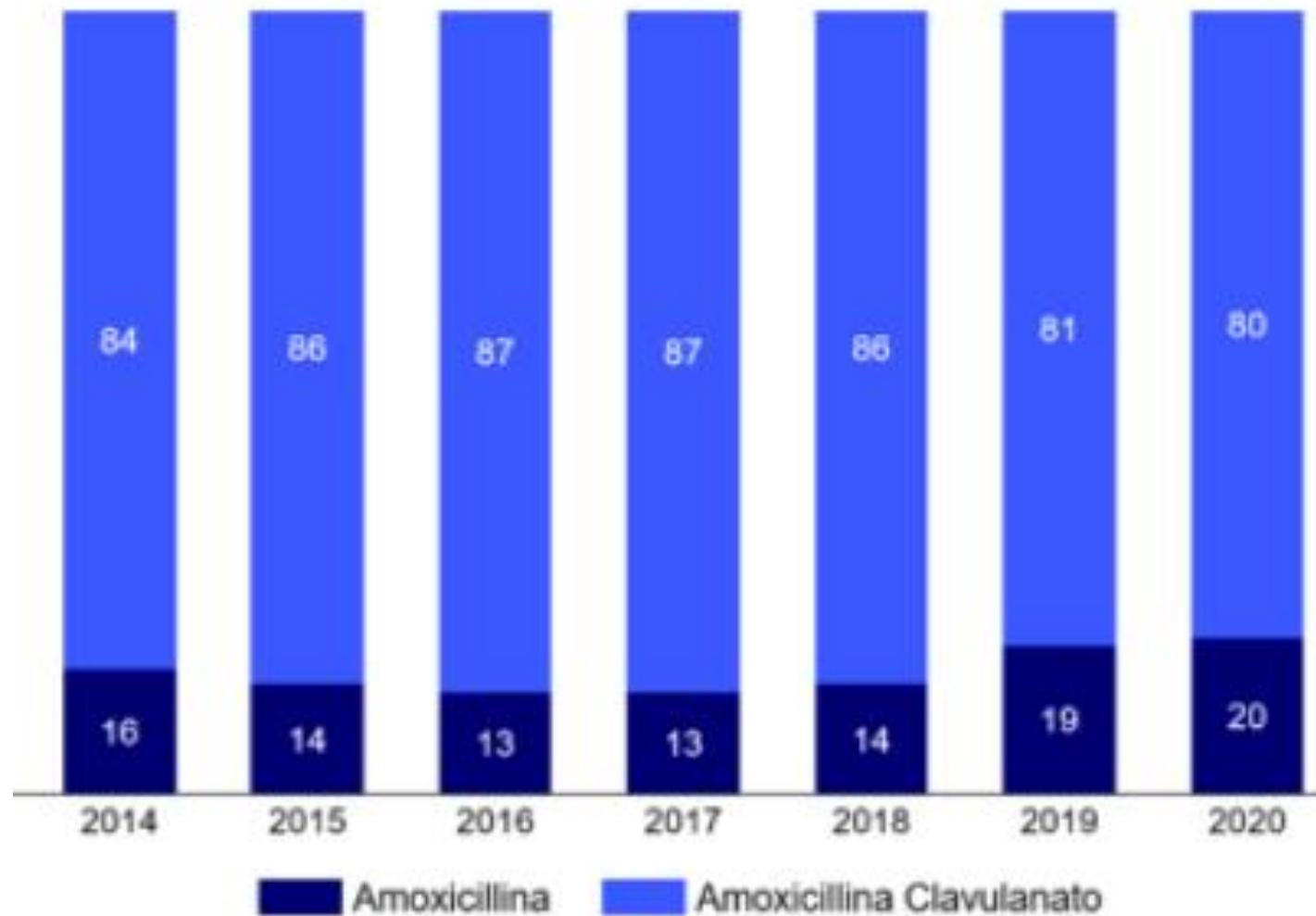


Consumo di antibiotici in Italia



Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Amoxicillina vs Amoxi/clav



L'utilizzo improprio di amoxicillina/acido clavulanico determina un ampliamento dello spettro di azione non necessario nella gran parte delle patologie infettive per le quali viene prescritta causando un aumento del rischio di fenomeni di antibiotico resistenza e reazione avverse attribuibili alla componente del clavulanato.

| | Amoxicillina | Amoxicillina - clavulanato |
|---------------|--------------|----------------------------|
| AV Centro | 28 | 72 |
| AV Nord-Ovest | 9 | 91 |
| AV Sud-Est | 20 | 80 |

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Piano Nazionale di Contrast all'Antibiotico-Resistenza PNCAR 2022-2025

Appendice: funghi, virus e parassiti



SORVEGLIANZA E MONITORAGGIO

- ABR
- ICA
- Uso antibiotici
- Monitoraggio ambientale



PREVENZIONE DELLE INFEZIONI

- ICA
- Malattie infettive e zoonosi



BUON USO ANTIBIOTICI

- Ambito umano
- Ambito veterinario
- Corretta gestione e smaltimento

Governance

Formazione

Informazione, comunicazione e trasparenza

Ricerca, innovazione e bioetica

Cooperazione nazionale e internazionale

Effects of national antibiotic stewardship and infection control strategies on hospital-associated and community-associated meticillin-resistant *Staphylococcus aureus* infections across a region of Scotland: a non-linear time-series study

| | Without intervention | With intervention | Marginal difference in MRSA prevalence density associated with successive interventions | | MRSA cases prevented per year (95% CI) |
|---|----------------------|-------------------|---|---------|--|
| | | | Absolute reduction (95% CI) | p value | Relative reduction (95% CI) |
| Hospitals | | | | | |
| Hand hygiene campaign (January, 2007) | 1.890 | 1.500 | 0.390 (-0.527 to 1.307) | 0.448 | 21% (-27 to 69) 246 (-316 to 822) |
| Universal screening (August, 2008) | 1.417 | 1.129 | 0.288 (-0.725 to 1.53) | 0.495 | 20% (-51 to 92) 180 (-444 to 796) |
| Hospital antibiotic stewardship (May, 2009) | 1.091 | 0.499 | 0.592 (0.001 to 1.180) | 0.049 | 54% (1 to 100) 355 (1 to 714) |
| Combined | 1.890 | 0.947 | 0.943 (0.267 to 1.619) | 0.006 | 50% (14 to 86) 592 (168 to 1017) |
| Community | | | | | |
| Indirect effects (hospital interventions)* | 0.071 | 0.045 | 0.026 (0.008 to 0.038) | 0.001 | 32% (11 to 54) 390 (128 to 652) |
| Primary care antibiotic stewardship (May, 2009) | 0.045 | 0.028 | 0.017 (0.004 to 0.029) | 0.012 | 37% (9 to 64) 281 (71 to 491) |
| Combined | 0.071 | 0.038 | 0.033 (0.018 to 0.048) | <0.0001 | 47% (25 to 68) 567 (311 to 822) |

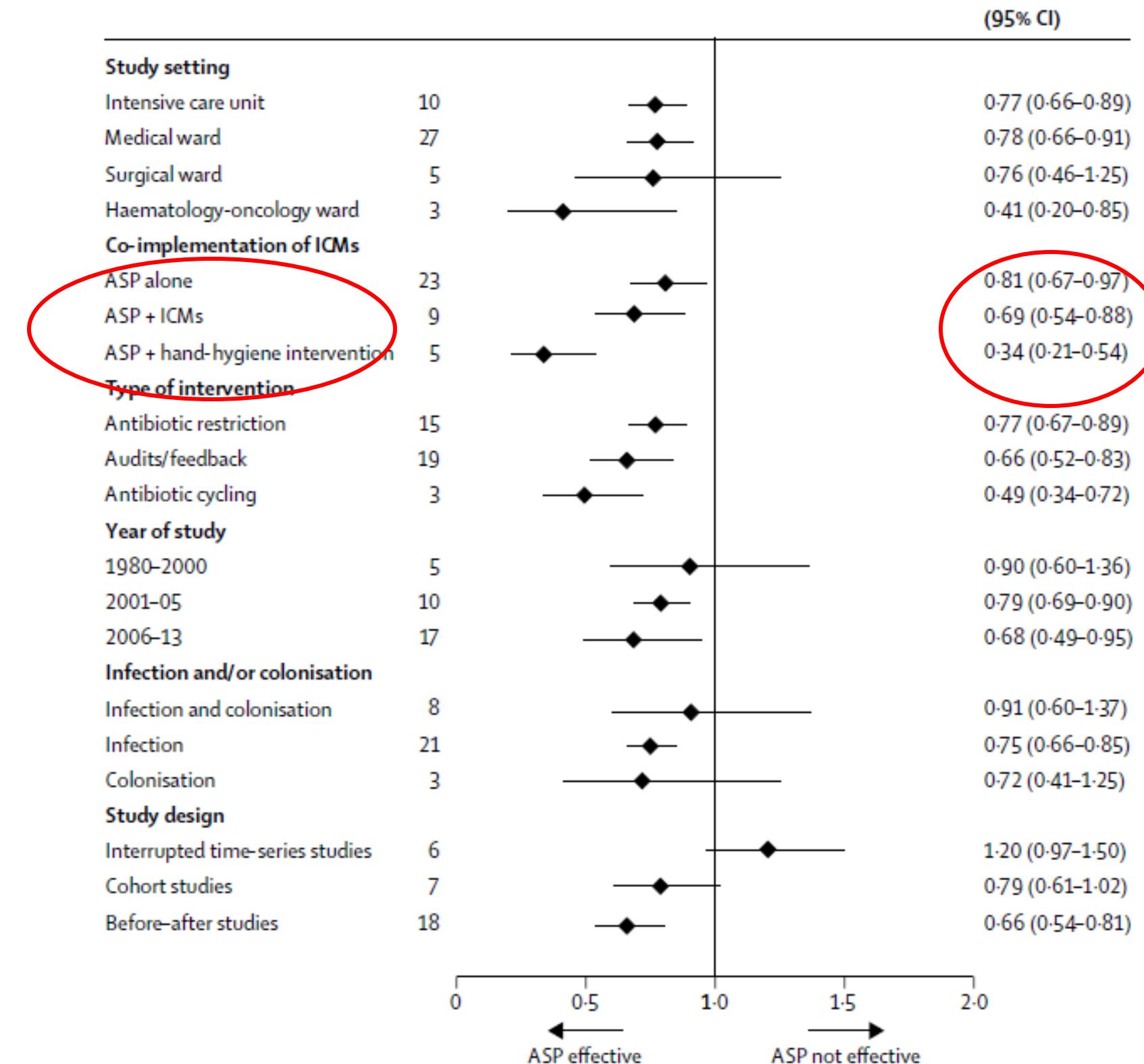
Lawes T et al, 2015

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Effect of antibiotic stewardship on the incidence of infection and colonisation with antibiotic-resistant bacteria and Clostridium difficile infection: a systematic review and meta-analysis

THE LANCET
Infectious Diseases

Baur D, et al. Lancet Infect Dis 2017



Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Infectious Diseases Society of America and the Society for Healthcare Epidemiology of America guidelines for developing an institutional program to enhance antimicrobial stewardship. Dellit TH, et al. Clin Infect Dis 2007; 44:159-177

Stewardship antimicobica :

Attività che ottimizza il trattamento antimicrobico e contempla scelta del farmaco, dose, via di somministrazione e durata della terapia

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

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Stewardship antimicrobica : obiettivo primario

Ottimizzare *l'outcome* clinico riducendo al minimo le possibili conseguenze :

- Tossicità
- Selezione di organismi patogeni (*C. difficile*)
- Insorgenza di resistenze

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

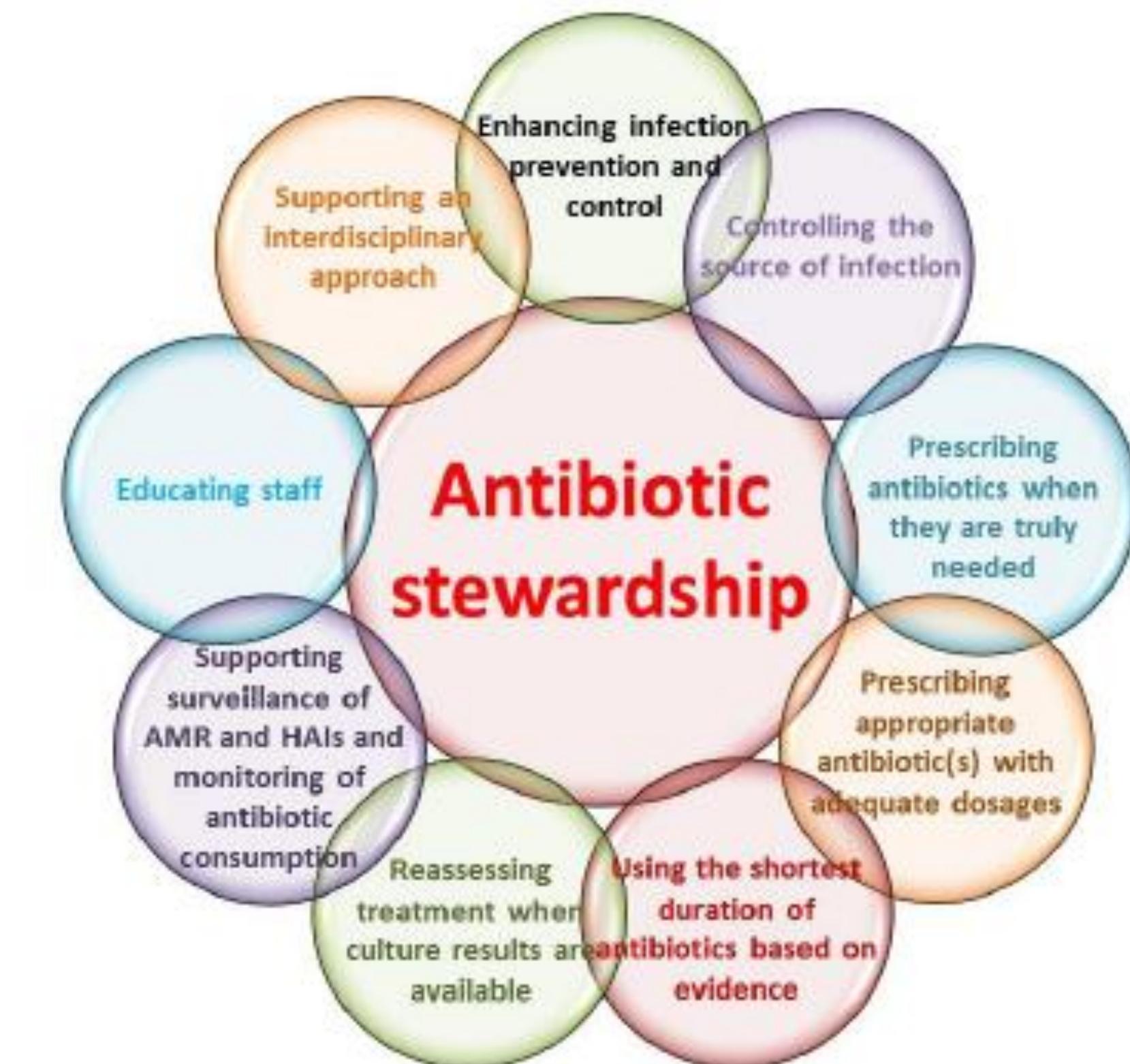
Infectious Diseases Society of America and the Society for Healthcare Epidemiology of America guidelines for developing an institutional program to enhance antimicrobial stewardship. Dellit TH, et al. Clin Infect Dis 2007; 44:159-177

Stewardship antimicrobica : obiettivo secondario

Ridurre i costi sanitari senza
inficiare la qualità della cura

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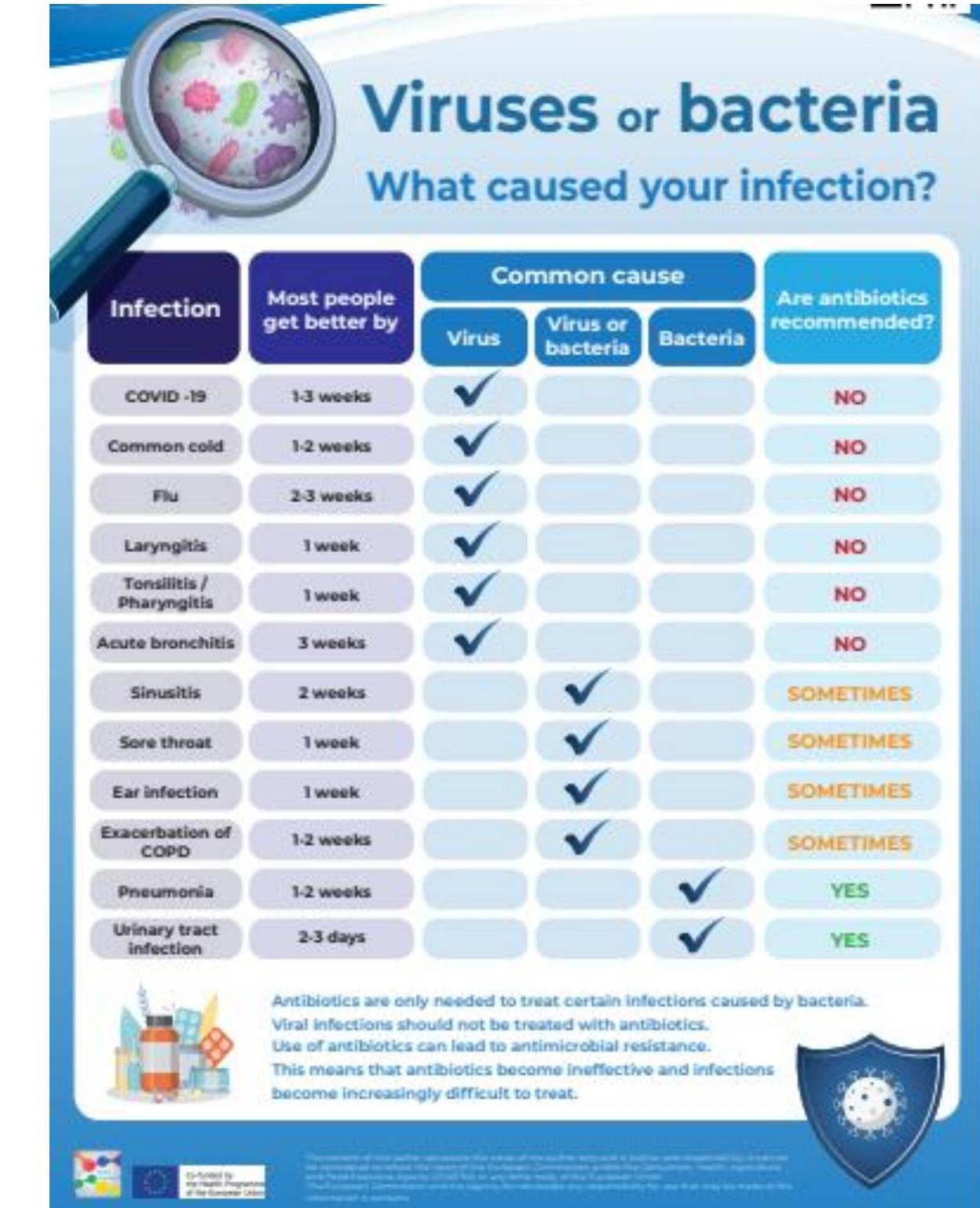
Dove è
possibile
agire?



Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico



Educazione...
Non solo gli operatori



Viruses or bacteria
What caused your infection?

| Infection | Most people get better by | Common cause | | | Are antibiotics recommended? |
|---------------------------|---------------------------|--------------|-------------------|----------|------------------------------|
| | | Virus | Virus or bacteria | Bacteria | |
| COVID-19 | 1-3 weeks | ✓ | | | NO |
| Common cold | 1-2 weeks | ✓ | | | NO |
| Flu | 2-3 weeks | ✓ | | | NO |
| Laryngitis | 1 week | ✓ | | | NO |
| Tonsillitis / Pharyngitis | 1 week | ✓ | | | NO |
| Acute bronchitis | 3 weeks | ✓ | | | NO |
| Sinusitis | 2 weeks | | ✓ | | SOMETIMES |
| Sore throat | 1 week | | ✓ | | SOMETIMES |
| Ear infection | 1 week | | ✓ | | SOMETIMES |
| Exacerbation of COPD | 1-2 weeks | | ✓ | | SOMETIMES |
| Pneumonia | 1-2 weeks | | ✓ | | YES |
| Urinary tract infection | 2-3 days | | ✓ | | YES |

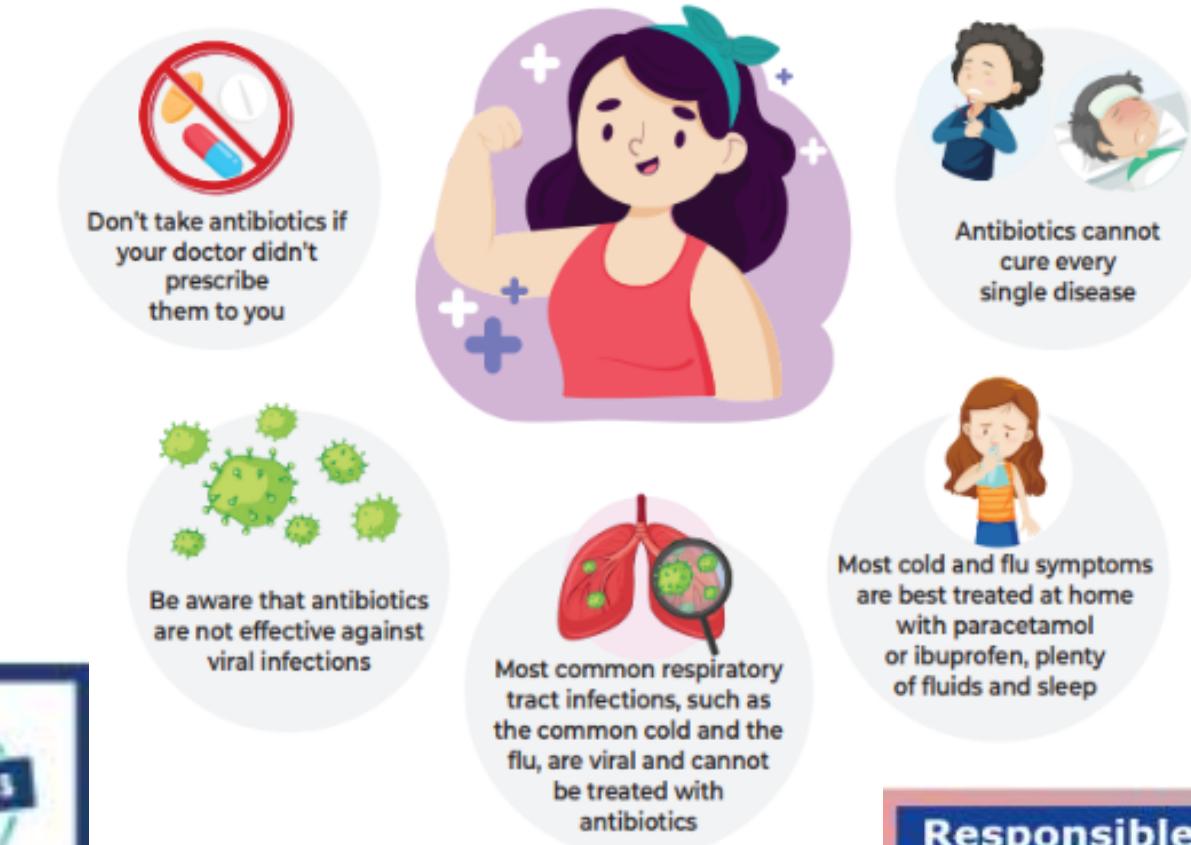
Antibiotics are only needed to treat certain infections caused by bacteria.
Viral infections should not be treated with antibiotics.
Use of antibiotics can lead to antimicrobial resistance.
This means that antibiotics become ineffective and infections become increasingly difficult to treat.

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico



HOW CAN YOU FIGHT IT?



ANTIBIOTIC RESISTANCE WHAT YOU CAN DO



Antibiotic resistance happens when bacteria change and become resistant to the antibiotics used to treat the infections they cause.

- Only use antibiotics when prescribed by a certified health professional
- Always take the full prescription, even if you feel better
- Never use left over antibiotics
- Never share antibiotics with others
- Prevent infections by regularly washing your hands, avoiding contact with sick people and keeping your vaccinations up to date

Responsible use of antibiotics: what's your role?



As a **patient** you should use antibiotics only when and as prescribed by your doctor.

Talk to your doctor or pharmacist for more information.

Never share antibiotics that were prescribed to you!

#EAAD #AMR #AntimicrobialResistance

EUROPEAN MEDICINES AGENCY

Interventi sulla terapia antibiotica

Regola delle «5 D»

- Right diagnosis
- Right drug
- Right dose (route of administration)
- Right duration
- (De-escalation)

Interventi sulla terapia antibiotica

Regola delle «5 D»

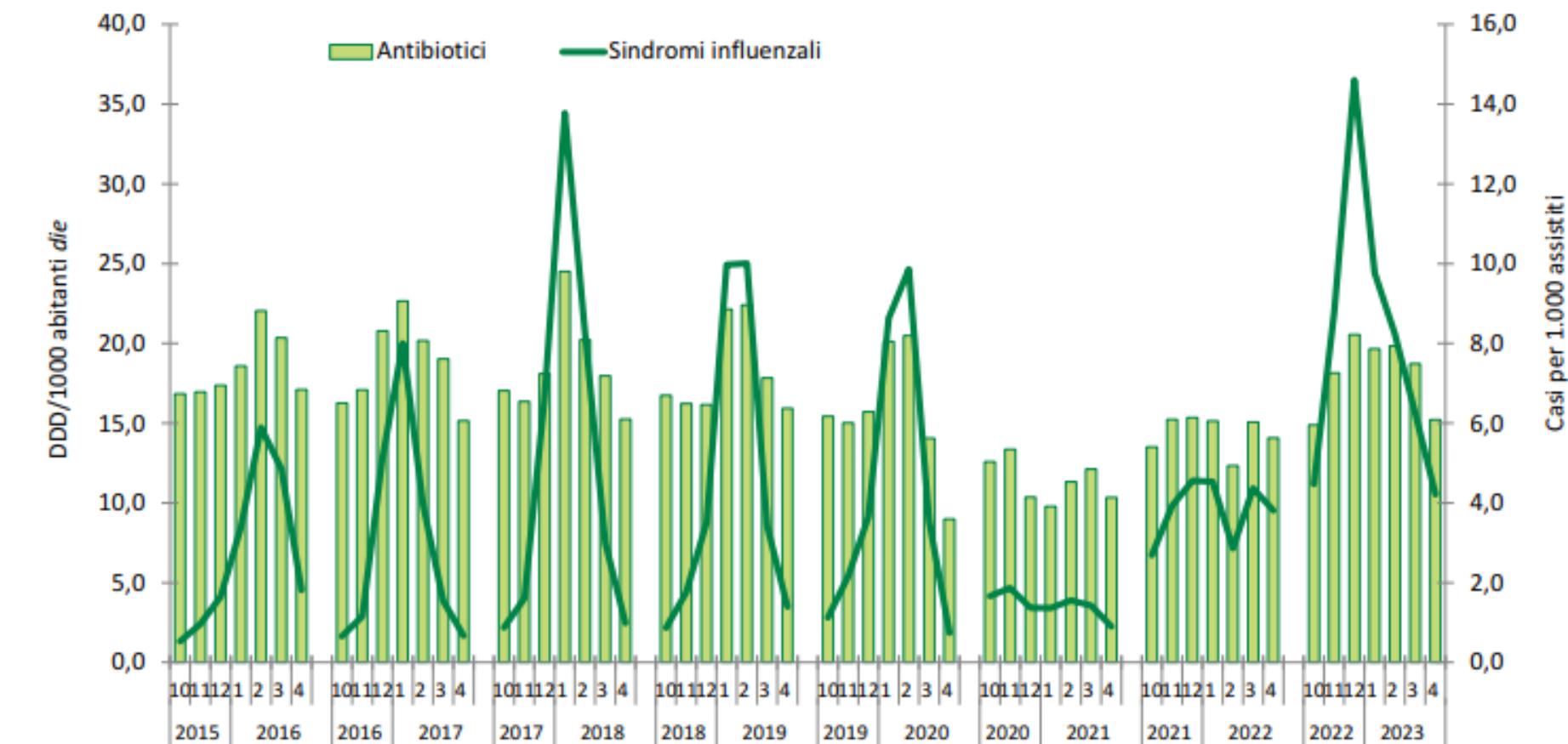
- **Right diagnosis**
- Right drug
- Right dose (route of administration)
- Right duration
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Trattare solo le vere
infezioni batteriche

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Interventi sulla terapia antibiotica

| Disease | Cases (n) | Antibiotic prescriptions (n) | Prescriptions (%) | Fever (%) |
|---------------------------------------|------------|------------------------------|-------------------|-------------|
| Rhinitis | 219 | 7 | 3.2 | 27.9 |
| Sinusitis | 21 | 5 | 23.8 | 23.8 |
| Non-group A streptococcal pharyngitis | 78 | 20 | 25.6 | 70.5 |
| Group A streptococcal pharyngitis | 32 | 32 | 100 | 65.6 |
| Otitis media | 102 | 74 | 72.5 | 46.1 |
| Laryngitis | 41 | 6 | 14.6 | 39 |
| Bronchitis | 167 | 93 | 55.7 | 67.3 |
| Bronchiolitis | 10 | 1 | 10 | 40 |
| Pneumonia | 8 | 8 | 100 | 37.5 |
| Total | 678 | 246 | 36.3 | 41.3 |

Picca M, 2023


Pediatric Infectious Diseases and Medical Microbiology



1. Don't routinely use antibiotics other than amoxicillin in the treatment of children with presumed community-acquired pneumonia (in the outpatient setting).
2. **Don't use a bag** for collection of **urine cultures** to diagnose urinary tract infections
3. **Don't routinely collect** or process specimens for **Clostridium difficile** testing in **infants less than one year** of age with diarrhea.
4. Don't routinely treat uncomplicated acute hematogenous osteomyelitis with prolonged intravenous therapy.

Raccomandazioni pediatriche

- 1. Non prescrivere** antibiotici nelle **patologie acute delle vie respiratorie** presumibilmente virali in età pediatrica (otiti, sinusiti, faringiti, bronchiti)
- 2. Non somministrare** terapia antibiotica ai bambini con **faringotonsillite acuta in assenza di conferma microbiologica** di infezione da Streptococcus pyogenes a seguito di test rapido o esame culturale.
- 3. Non fare diagnosi di Infezione delle Vie Urinarie** in base al solo **esame culturale** delle urine.
- 4. Non eseguire la profilassi antibiotica** peri-operatoria quando questa non sia **necessaria**.
- 5. Non superare le 24 ore di profilassi antibiotica** dopo la chirurgia, la durata della profilassi dovrebbe essere la più breve possibile.

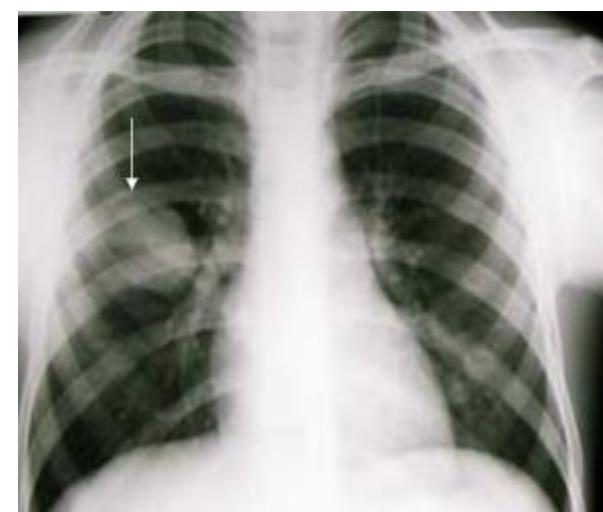
Interventi sulla terapia antibiotica

Regola delle «5 D»

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Principali infezioni in età pediatrica

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico



Management of Acute Pharyngitis in Children: Summary of the Italian National Institute of Health Guidelines

Amoxicillina

Updated Guidelines for the Management of Acute Otitis Media in Children by the Italian Society of Pediatrics

NON COMPLICATE
Amoxicillina

Marchisio P, Galli L, Chiappini E Pediatr Infect Dis J. 2019

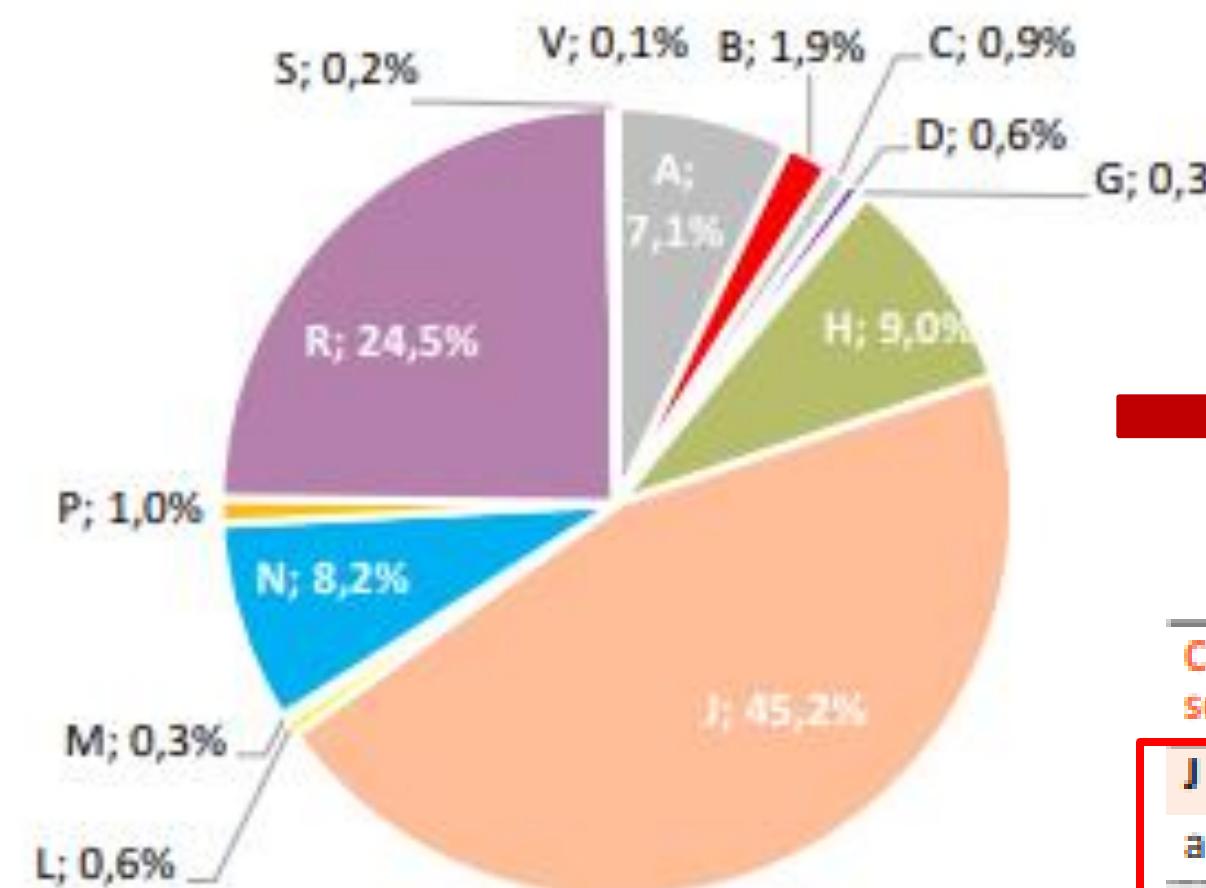
British Thoracic Society guidelines for the management of community acquired pneumonia in children: update 2011

Amoxicillina

The Management of Community-Acquired Pneumonia in Infants and Children Older Than 3 Months of Age: Clinical Practice Guidelines by the Pediatric Infectious Diseases Society and the Infectious Diseases Society of America 



Utilizzo dei farmaci in età pediatrica



| Categoria terapeutica/ sostanza | Confezioni (per 1000 bambini) | Δ % 23-22 | Δ % 22-21 | Prevalenza (per 1000 bambini) | Rapp M/F |
|---|----------------------------------|--------------|--------------|----------------------------------|-------------|
| J - Antinfettivi per uso sistemico | 977,3 | 29,9 | 53,3 | 482,0 | 1,1 |
| amoxicillina/acido clavulanico | 409,7 | 45,4 | 57,3 | 210,4 | 1,1 |
| amoxicillina | 149,8 | 27,7 | 67,6 | 72,5 | 1,1 |
| cefixima | 123,3 | 36,1 | 69,7 | 79,3 | 1,1 |
| azitromicina | 100,2 | 8,7 | 28,9 | 66,2 | 1,1 |

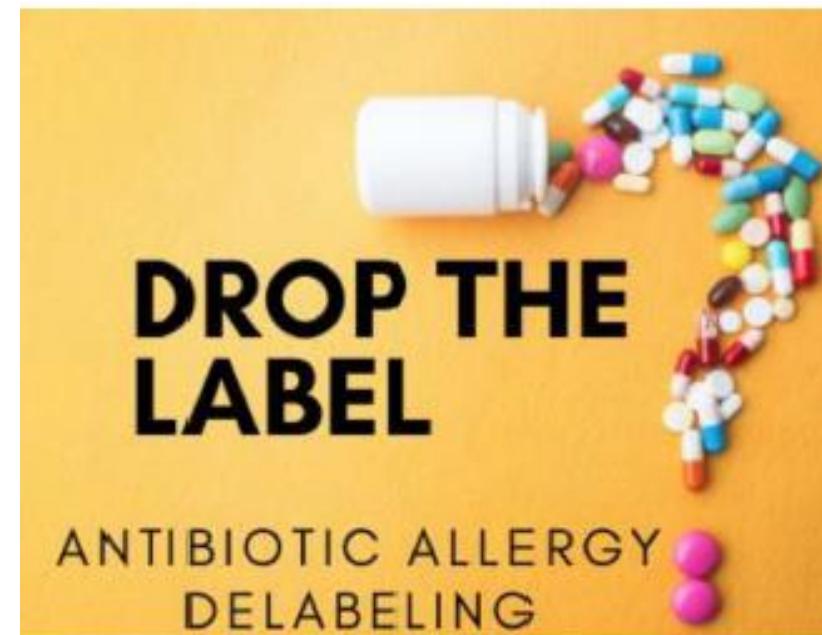
The Impact of Clinical Pathways on Antibiotic Prescribing for Acute Otitis Media and Pharyngitis in the Emergency Department

| Patients included | ACUTE OTITIS MEDIA (AOM) | | | | p value | GAS PHARYNGITIS | | | | |
|--|--------------------------|------|--------------------------|------|---------|-------------------------|------|--------------------------|------|---------|
| | Pre-intervention period | | Post-intervention period | | | Pre-intervention period | | Post-intervention period | | |
| | n | % | n | % | | n | % | n | % | |
| TREATMENT | | | | | | | | | | |
| “Wait and see” for AOM or no antibiotic treatment for GAS pharyngitis | 64 | 21.7 | 92 | 33.1 | p<0.01 | 147 | 49.3 | 200 | 54.6 | p=0.17 |
| Antibiotic therapy | 231 | 78.3 | 186 | 66.9 | p<0.01 | 151 | 50.7 | 166 | 45.4 | p=0.17 |
| TYPE OF ANTIBIOTICS | | | | | | | | | | |
| Amoxicillin | 74 | 32.0 | 96 | 51.6 | p<0.001 | 81 | 53.6 | 155 | 93.4 | p<0.001 |
| Broad spectrum (amoxi-clavulanate +cephalosporins+ macrolides) | 157 | 68.0 | 90 | 48.4 | p<0.001 | 70 | 46.4 | 11 | 6.6 | p<0.001 |
| Amoxicillin-clavulanate | 106 | 45.9 | 70 | 37.6 | p=0.09 | 60 | 39.7 | 5 | 3.0 | p<0.001 |
| Cephalosporins | 47 | 20.3 | 16 | 8.6 | p<0.001 | 10 | 6.6 | 6 | 3.6 | p=0.28 |
| Macrolides | 4 | 1.7 | 4 | 2.2 | p=0.76 | - | - | - | - | - |

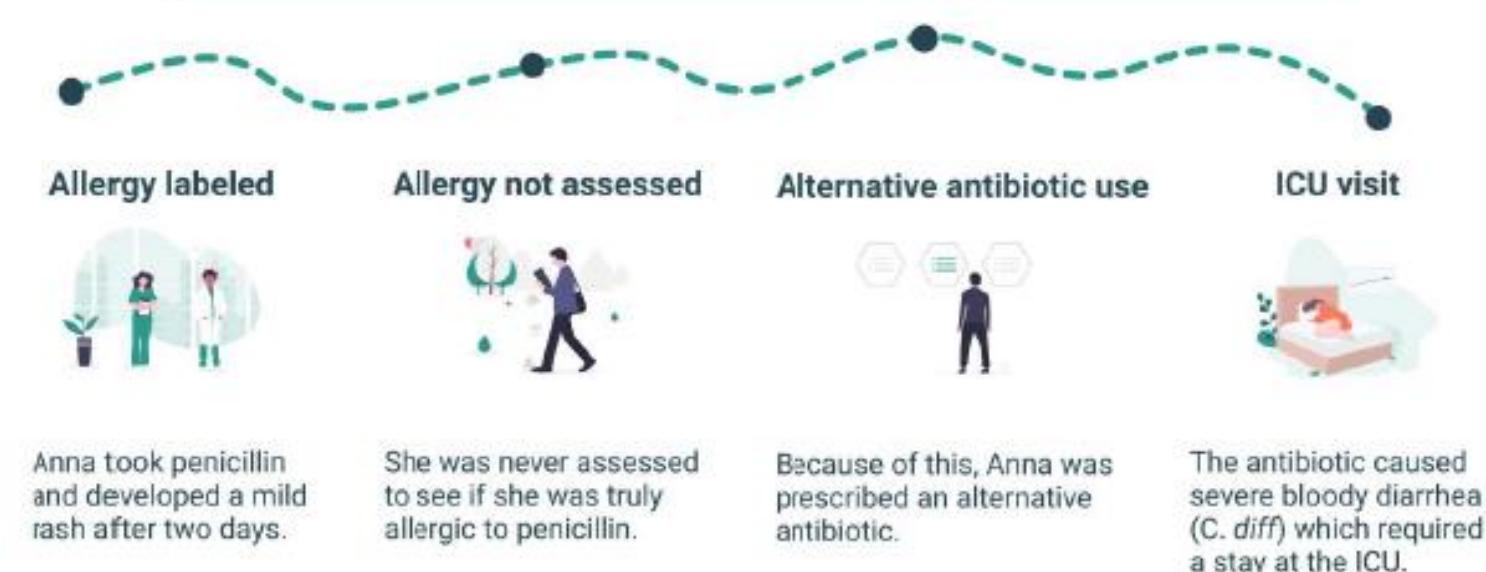
Donà D, et al. 2018

Il de-labeling dell'allergia ai beta-lattamici

Because most **antibiotic allergy labels** acquired in **childhood** are carried into adulthood, the overlabeling of antibiotic allergy is a liability that leads to unnecessary long-term **health care risks, costs, and antibiotic resistance**.



Harms of false penicillin allergy labels: *Meet Anna*



Anna was never allergic to penicillin, her ICU visit could have been prevented.

<https://www.dropthelabel.ca>

Norton AE, et al. Pediatrics 2018

Interventi sulla terapia antibiotica

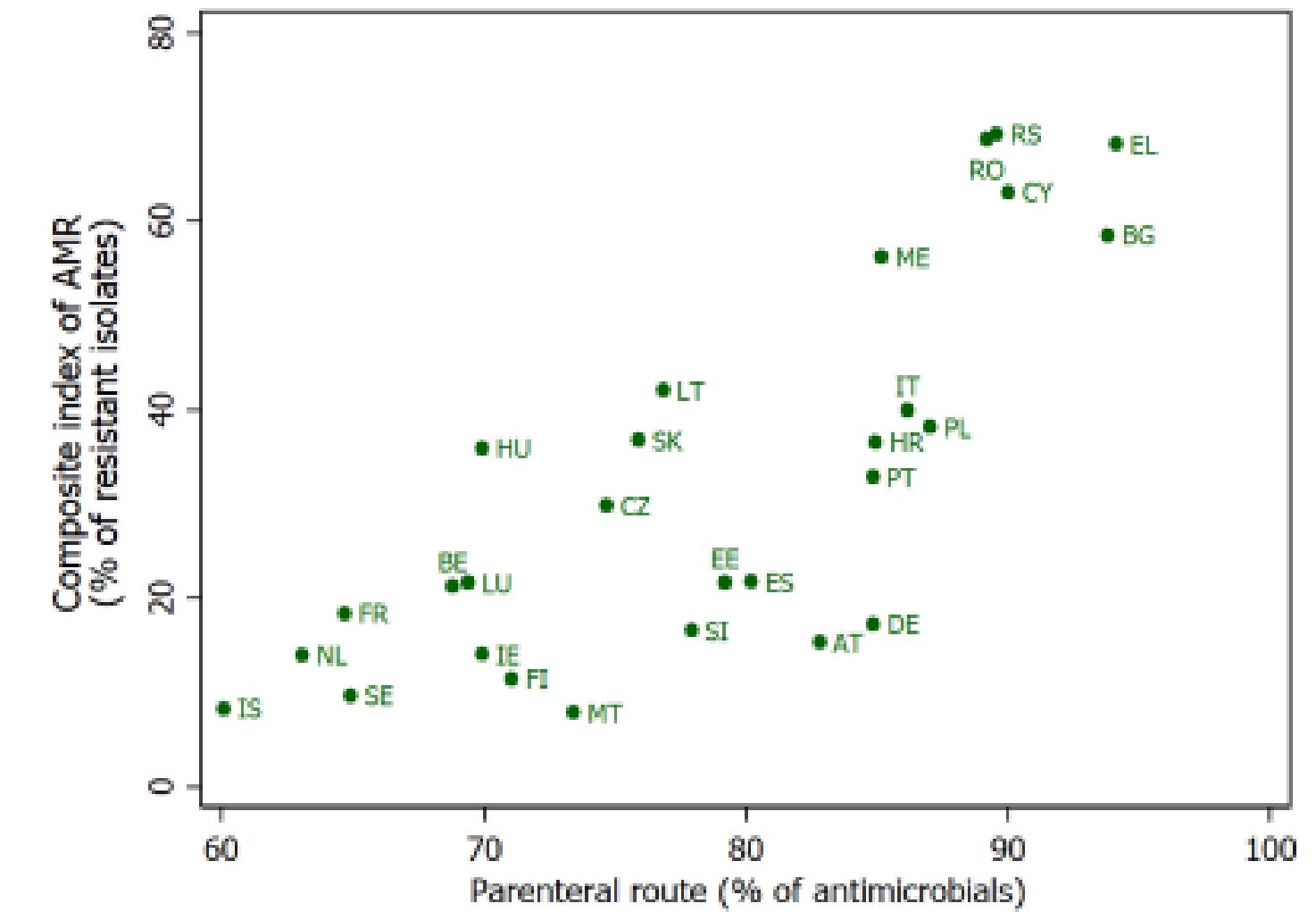
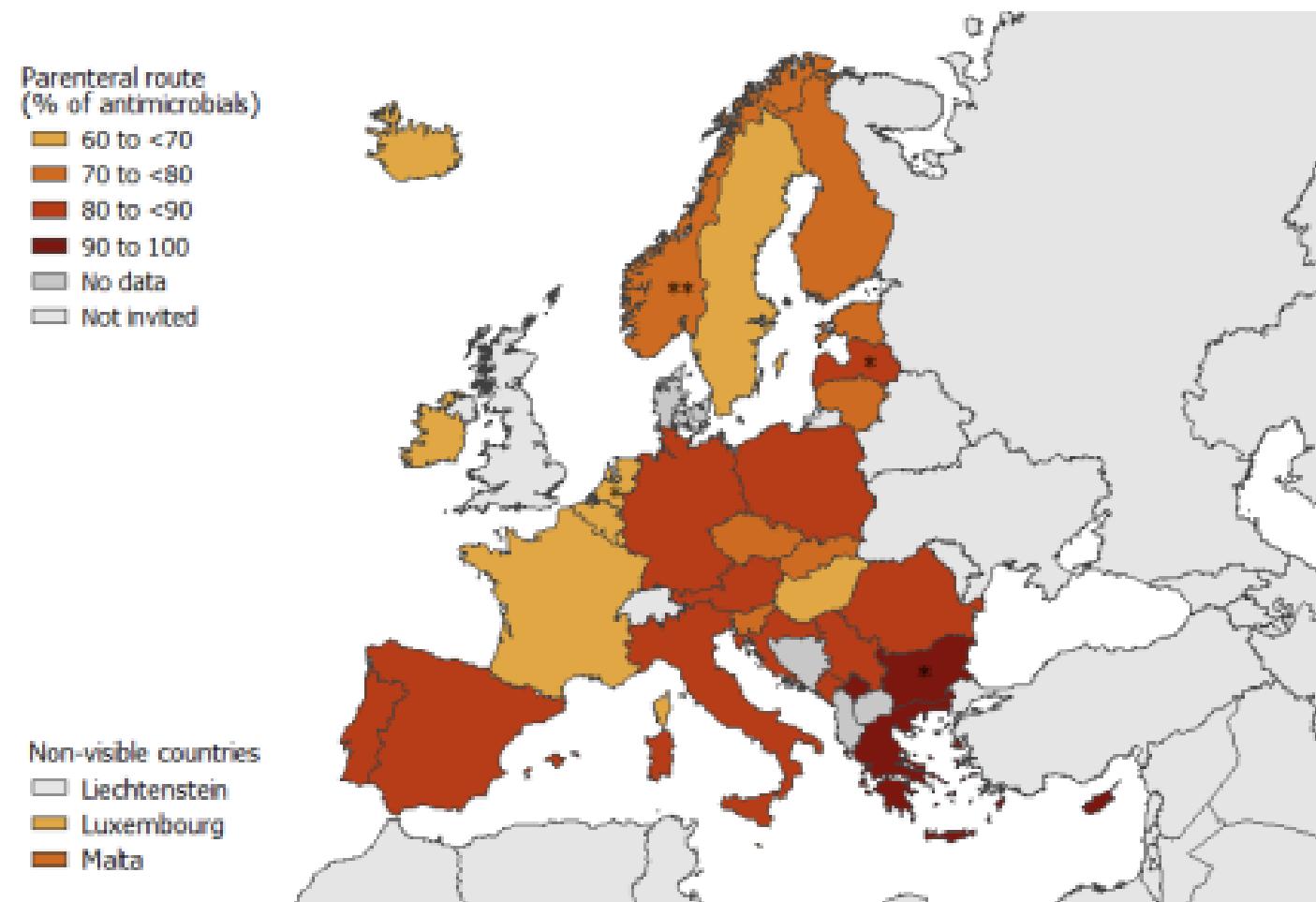
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La % di antibiotici prescritti per via parenterale correla fortemente con il composite index of AMR ($p > 0.001$)

Figure 61. Percentage of antimicrobials for which the route of administration was parenteral, ECDC PPS 2022–2023



ECDC, 2023

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

Antibiotici adatti allo switch per os (biodisponibilità > 90%)

Amoxi-clav, Cefalexina, Clindamicina, TMP-SMX, metronidazolo, fluconazolo



Rivalutare dopo 48 ora dall'inizio della terapia antibiotica

- Clinicamente stabile (apiretico, indici di flogosi in calo, parametri vitali stabili)
 - Capace di assumere farmaci per os

Si



No

**Rivaluta ogni
24 ore**

**Passaggio a
terapia orale**

Interventi sulla terapia antibiotica

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Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

PEDIATRICS Volume 153, number 1, January 2024:e2023062598

Effectiveness and safety of modified fully oral 9-month treatment regimens for rifampicin-resistant tuberculosis: a prospective cohort study



Lancet Infect Dis 2024;
24: 1151-61

JAMA Pediatrics | Original Investigation

Short-Course Therapy for Urinary Tract Infections in Children
The SCOUT Randomized Clinical Trial

JAMA Pediatr. 2023;177(8):782-789.

Theoklis Zaoutis, MD; Nader Shaikh, MD; Brian T. Fisher, DO; Susan E. Coffin, MD; Sonika Bhatnagar, MD; Kevin J. Downes, MD; Jeffrey S. Gerber, MD; Timothy R. Shope, MD; Judith M. Martin, MD; Gysella B. Muniz, MD; Michael Green, MD; Jennifer P. Nagg, RN; Sage R. Myers, MD; Rakesh D. Mistry, MD; Shawn O'Connor, BS; Walter Faig, PhD; Stephen Black, MS; Elizabeth Rowley, PhD; Kellie Liston, BA; Alejandro Hoberman, MD

JAMA Pediatrics | Original Investigation

Short-Course vs Long-Course Antibiotic Therapy for Children With Nonsevere Community-Acquired Pneumonia
A Systematic Review and Meta-analysis

JAMA Pediatr. 2022;176(12):1199-120

Short Oral Antibiotic Therapy for Pediatric Febrile Urinary Tract Infections: A Randomized Trial

Giovanni Montini, MD,^{a,b} Antimo Tessitore, MD,^c Karen Console, MD,^c Luca Ronfani, MD,^d Egidio Barbi, MD,^{a,c} Marco Pennesi, MD,^e on behalf of the STOP Trial Group



ESTABLISHED IN 1812

MARCH 10, 2022

VOL. 386 NO. 10

Shorter Treatment for Nonsevere Tuberculosis in African and Indian Children

A. Turkova, G.H. Wills, E. Wobudeya, C. Chabala, M. Palmer, A. Kinikar, S. Hissar, L. Choo, P. Musoke, V. Mulenga, V. Mave, B. Joseph, K. LeBeau, M.J. Thomason, R.B. Mboizi, M. Kapasa, M.M. van der Zalm, P. Raichur, P.K. Bhavani, H. McIlheran, A.-M. Demers, R. Aarnoutse, J. Love-Koh, J.A. Seddon, S.B. Welch, S.M. Graham, A.C. Hesseling, D.M. Gibb, and A.M. Crook, for the SHINE Trial Team*

Pediatric Infectious Diseases and Medical Microbiology



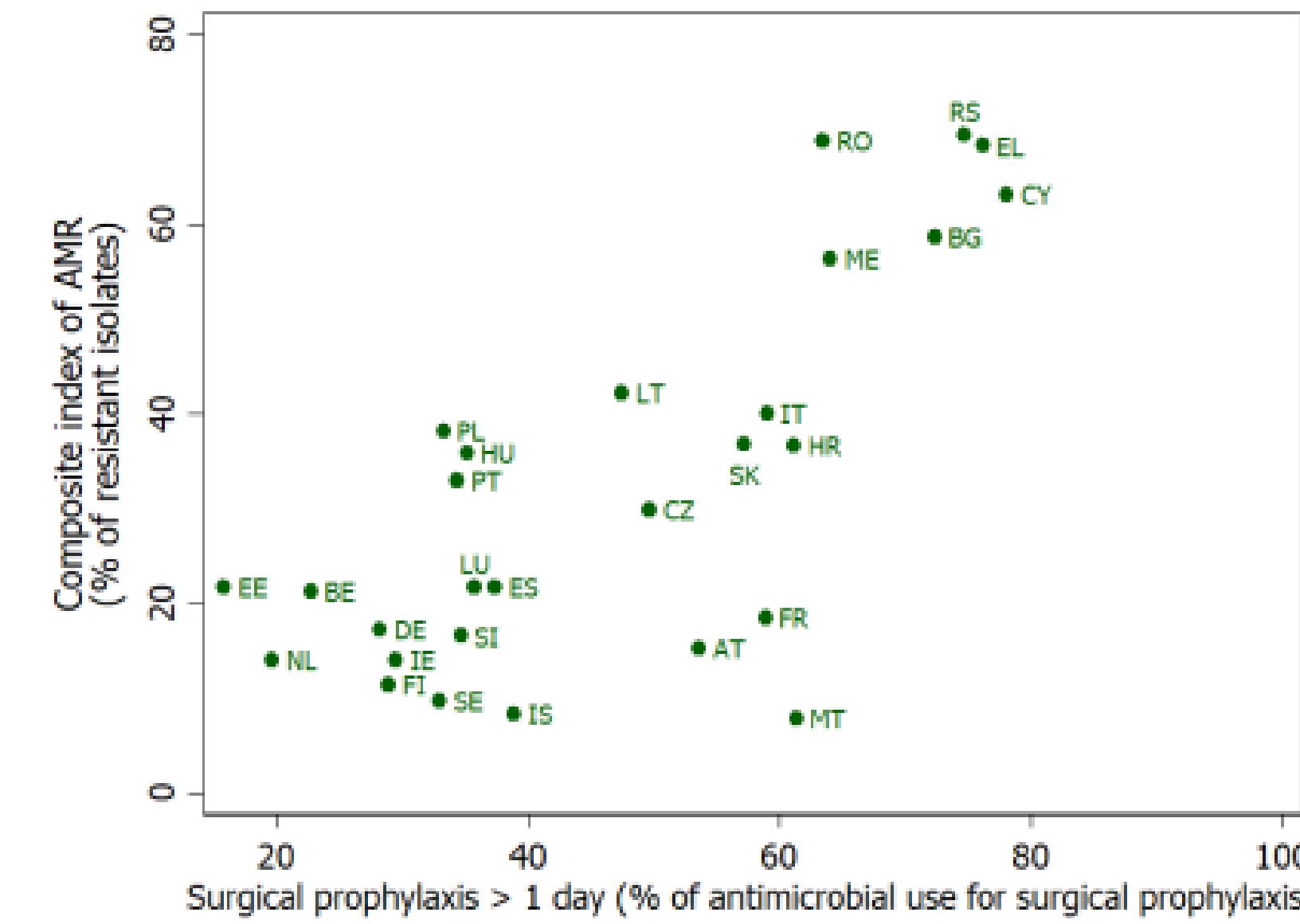
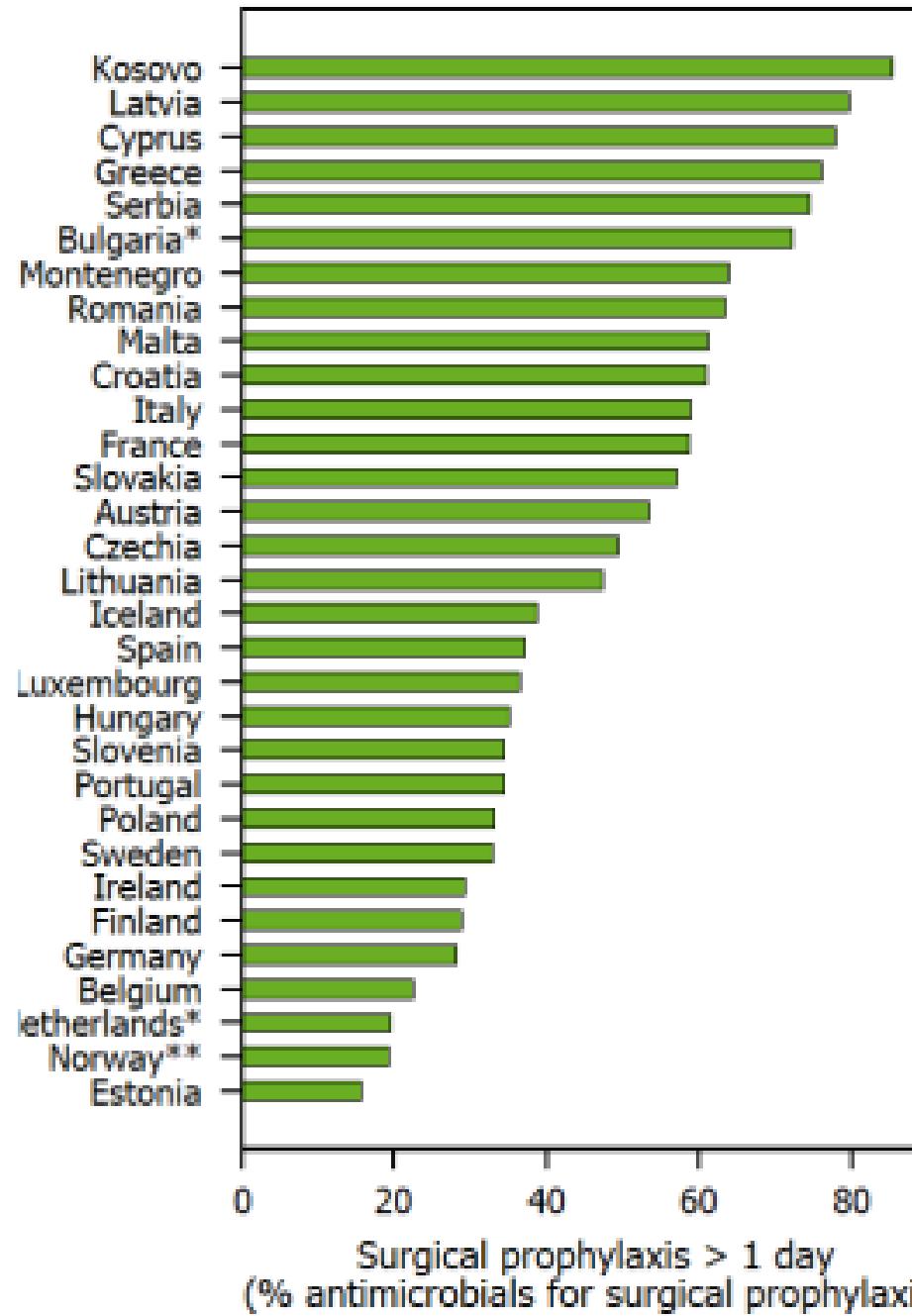
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- 3. Non fare diagnosi di Infezione delle Vie Urinarie** in base al solo **esame culturale** delle urine.
- 4. Non eseguire la profilassi antibiotica** peri-operatoria quando questa non sia **necessaria**.
- 5. Non superare le 24 ore di profilassi antibiotica** dopo la chirurgia, la durata della profilassi dovrebbe essere la più breve possibile.

Strategie innovative per la prevenzione delle infezioni correlate all'assistenza in ambito pediatrico

La % di profilassi perioperatoria > 1 giorno correla fortemente con il composite index of AMR ($p > 0.001$)

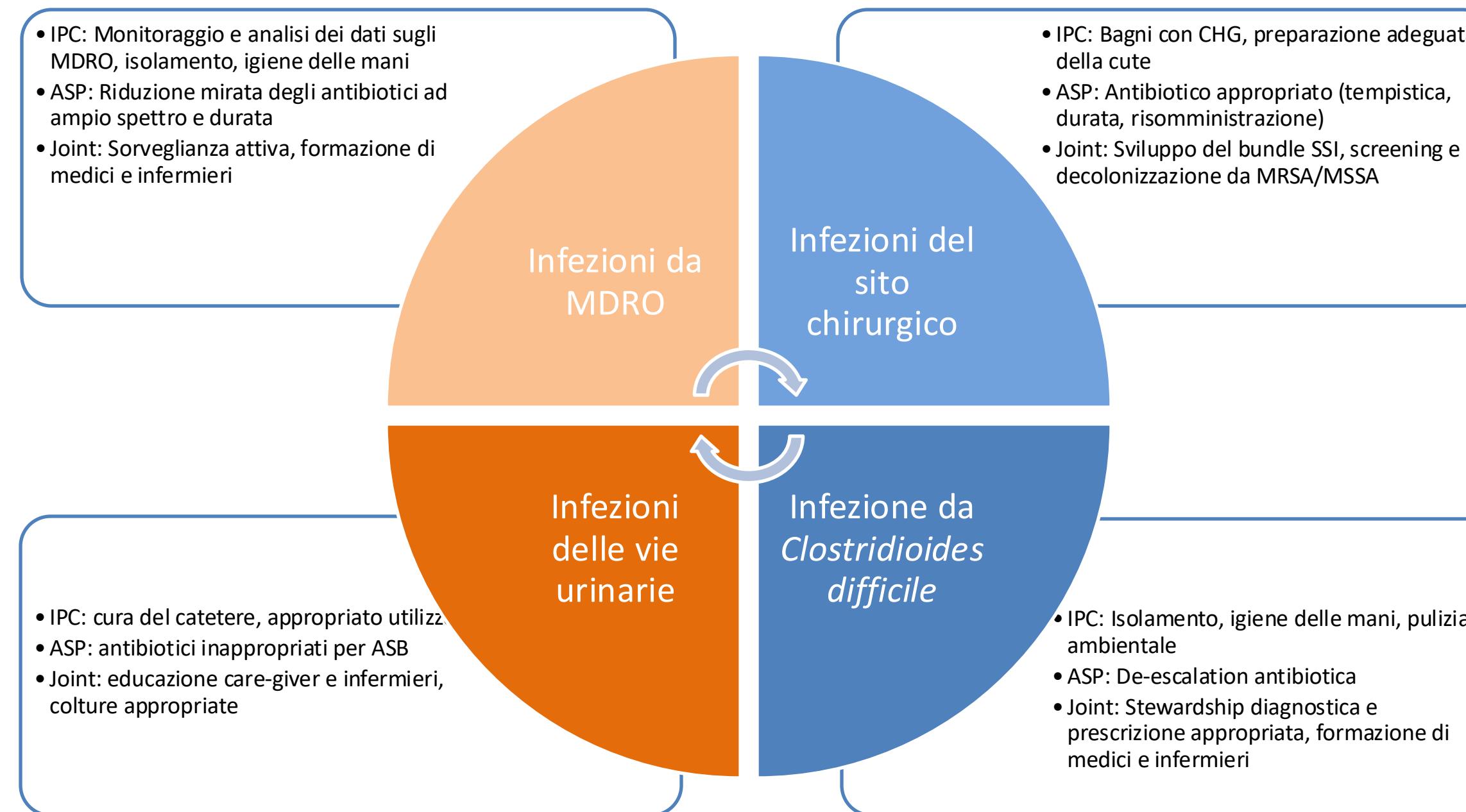


ECDC, 2023

Cosa facciamo noi?

- Monitoraggio consumo di antibiotici in termini di DDD e DOT
- Monitoraggio dei microrganismi multiresistenti
- Monitoraggio del Clostridioides difficile
- Sistema di pre-autorizzazione (carbapenemi, teicoplanina e tutti i farmaci Reserve)
- Rivalutazione delle prescrizioni antibiotiche (appropriatezza, dose, interazioni, passaggio dalla terapia endovenosa a orale e durata)
- Redazione, diffusione e monitoraggio protocolli sulla terapia antibiotica (sepsi, IVU, Osteomieliti/Artriti, polmoniti, faringiti, profilassi perioperatoria)
- Formazione degli operatori
- Educazione di pazienti e famiglie

Dove agire assieme all'IPC?



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