

Antimicrobial Stewardship Metrics What to do?

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Disclosure

- I have no actual or potential conflict of interest in relation to this program or presentation
- Commercial Interests:
 - none

*There are no standard,
universally accepted metrics,*

*.....for assessing the effectiveness of
antimicrobial stewardship programs
(ASPs)*

Akpan. *Antibiotics* 2016;5:5. Dodds Ashley. *Clin Infect Dis* 2014 59(s3:112). Nagel. *Clin Infect Dis* 2014;59(S3):S146.

Goals of ASPs

.....optimal clinical outcomes

.....appropriate use

*.....limit the selection of antimicrobial
resistant strains*

*ASPs are struggling to identify
appropriate measures of success;*

Nevertheless.....

*Accurate measurement is required
before any improvement can take
place*

Morris. *Current Treatment Options in Infectious Diseases* 2014; 6:101.

Reddy. *Expert Rev Anti Infect Ther*; 2015; 13:843

Step-back....

Can we learn from other programs that have demonstrated the value of their efforts?¹

Infection Control, in 2016....

...interventions that prevent bloodstream infections can be of high value...specifically;

- 57% fewer bloodstream infections
- Net savings: \$1.85 million/hospital over 3 years²

¹Schwartz. *Clin Infect Dis* 2016; 63:450. ²Nuckols. Economic evaluation of quality improvement interventions for bloodstream infections related to central catheters. A systematic review. *JAMA Intern Med.* 2016;176(12):1843.

Objectives

1. Review milestones in the emergence of Infection Control Programs (ICPs); focusing on the development of metrics for ICPs.
2. Review Joint Commission's expectations for ASPs with respect to metrics.
3. Review for ASPs:
 - Commonly recommended process and outcome metrics
 - Selecting process measures and performance metrics to measure the performance of ASP's interventions.
4. Review knowledge gaps in the area of ASP metrics.

Infection Surveillance and Control Programs (ISCPs)

50+ years... and still evolving....

Infection Surveillance and Control Programs (ISCPs)

- 1950s – 60s
 - *Staphylococcal* pandemic (pcn- resistant *S. aureus*)
“Antibiotic resistance became a serious problem.”
 - Voluntary formation of Infection Control Programs
- 1960s
 - CDC recommends regular surveillance
 - Data collection to inform rational infection control measures

1960s-70s

Who will fund these programs?

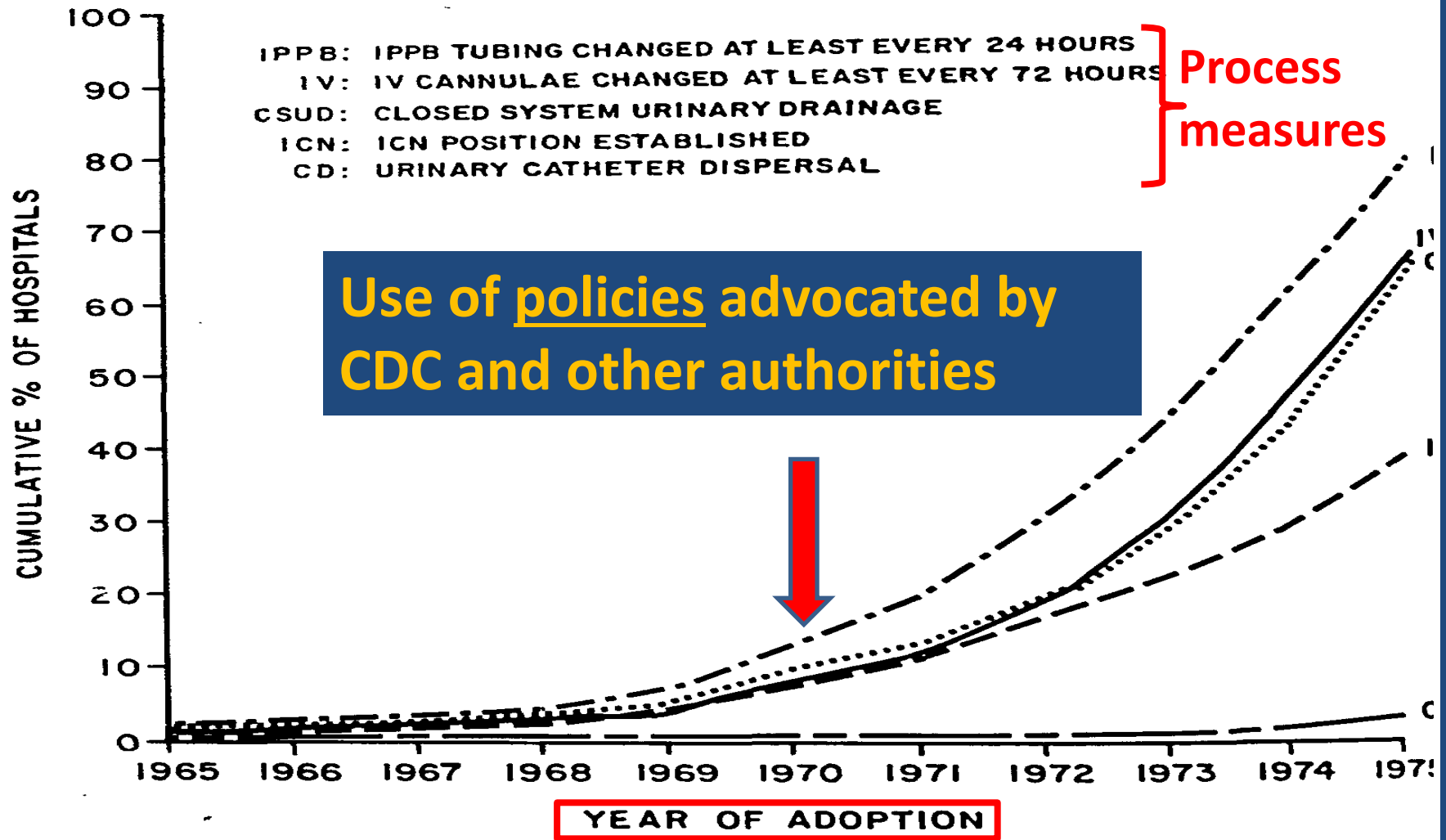
Inadequate evidence to
mandate infection control
programs

Study on the Effectiveness of Nosocomial Infection Control (SENIC) Project^{1,2} (CDC 1970-76)

Infection control program	<u>Outcome</u> Hospital infection rate
Effective ³	32% reduction
Not effective	18% increase

¹Haley. *Am J Epidemiol* 1985; 121:182. ²Dixon. *MMWR* 2011; 60:58. ³“Control index” measures the intensity of efforts to intervene to reduce infection risk. Effective program elements: trained personnel, surveillance, active preventive interventions, regular reporting surgical wound infection rates to surgeons.

THE EMERGENCE OF INFECTION SURVEILLANCE AND CONTROL PROGRAMS IN US HOSPITALS: AN ASSESSMENT, 1976



1976

Joint Commission Standard for
Infection Surveillance & Control
Programs

2017

Joint Commission Standard for
Antimicrobial Stewardship
Programs

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Joint Commission 2017

What type of antimicrobial stewardship data should organizations collect, analyze, and report?

....not requiring any specific antimicrobial stewardship data

.....the organization must determine the antimicrobial stewardship data it will collect, analyze, and report

Where to start.....performance metrics

Good faith effort to foster real quality improvement

Efforts should not.....

.....create incentives for providers to
improve measured performance,

...without truly improving quality of care

”Not just a “checkbox”

Performance metrics - *Imperatives for ASPs*

Focus on an area of an ASP that is:

- Important/problematic to the organization
- Marked variation in practice
- Good evidence to support a practice which can be simply measured

Performance metrics – case study

- Piperacillin/tazobactam plus vancomycin
 - associated with a higher risk for nephrotoxicity
 - compared to alternative beta-lactams plus vancomycin^{1,2}
- Commonly used antibiotic regimen

¹Navalkele. *Clin Infect Dis* 2017; 64:116. ²Hammond. *Clin Infect Dis* 2017; 64:666.

Steps to Quality Improvement^{1,2,3}

WHAT are we trying to accomplish?

Avoid PT-V, promote Cef-V, +/- metro⁴

WHY is it important?

Nephrotoxicity risk

WHO is the specific target population?

Adult ED and hospitalized patients

WHEN will this be carried out?

HOW will this be carried out?

WHAT is the data source?

¹McGowan. *J Antimicrob Chemother* 2016; 71:2370. ²Rubin. *Int J Quality in Health Care* 2001; 6:489. ³McGlynn. *Am J Prev Med* 1998; 14(3S):14. ⁴PT-V (piperacillin/tazobactam plus vancomycin; Cef-V, +/- metro (cefepime plus vancomycin, +/- metronidazole)

Steps to Quality Improvement (cont)

- WHAT are our measurable goals?

	Pre-intervention	Post-intervention
% cases on PT-V		
% cases on Cefepime-V, +/- metronidazole		

- HOW will we know our changes are working?

- Less nephrotoxicity
- Requires risk-adjustment for nephrotoxicity

Steps to Quality Improvement (cont)

- HOW do we know if the intervention is harmful?

(Measure unintended consequences)

- Omitting metronidazole when indicated e.g., intra-abdominal infections (cefepime-based)
- Change in *C. difficile* incidence
- Rates of super-infection due to *Enterococcus*, ESBL-producing gram-negatives

Unintended consequences (cont)

Intervention	Unintended consequences
Targeting specific agents	Increase in use of other agents, especially agents with a similar spectrum
e.g., reduce piperacillin/tazobactam	Increase use of other anti-pseudomonal agents (e.g., carbapenems)
Surgical prophylaxis ¹	Increased gentamicin nephrotoxicity
Replace cephalosporins with gentamicin	

¹Weeraporn. Ann Transl Med 2017; 5:100.

Steps to Quality Improvement (cont)

HOW completely was the intervention implemented?

% implemented =

$$\frac{\text{\# cases assessed}}{\text{Total \# of cases}} \times 100$$

Validity of data - **alternate reasons for results**

Example: Intervention reduces
specific antibiotic usage

- Intervention
- Shorter length of stay
- Fewer admissions
- Switch to other antibiotics
 - Assessing overall antibiotic use can help sort this out

**Reason
for
reduction
in usage?**

*Your experience with a Joint
Commission survey of your
ASP?*

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Quality Improvement

Defining key processes for improvement and metrics to monitor performance

Processes

- Interventions
- Quality indicators

**Demonstrated
linkage to
improve
outcome**

Outcomes

Resources: Process and Outcome Metrics

Hospitals

- Core Elements¹
- National Quality Partners Playbook on Antibiotic Stewardship²
- Systematic review of Quality Indicators³

Nursing Homes

- Core Elements⁴

Outpatient

- Core Elements⁵

¹ <https://www.cdc.gov/getsmart/healthcare/pdfs/core-elements.pdf>.

² [http://www.qualityforum.org/Publications/2016/05/National Quality_Partners_Playbook__Antibiotic_Stewardship_in_Acute_Care.aspx](http://www.qualityforum.org/Publications/2016/05/National_Quality_Partners_Playbook__Antibiotic_Stewardship_in_Acute_Care.aspx).

³<https://www.cdc.gov/longtermcare/prevention/antibiotic-stewardship.html>. ⁴WP1A quality indicators and quantity metrics of antibiotic use. 2016. Available from: http://driveab.eu/wp-content/uploads/2014/09/WP1A_Final-QMs-QIs_final.pdf. ⁵<https://www.cdc.gov/mmwr/volumes/65/rr/rr6506a1.htm>

Infection-specific quality indicators

- Pneumonia^{1,2}
- Sepsis³
- Urinary tract⁴
- Appropriate antibiotic use
in hospitalized adults⁵

Expert consensus

- Guideline adherence
- Obtain cultures
- Targeted therapy
- Others

¹Nathwani. *Clin Infect Dis* 2002; 34:318. ²Schouten. *Clin Infect Dis* 2005; 41:450. ³van den Bosch. *BMC Infect Dis* 2014; 14:345. ⁴Spoorenberg. *Clin Infect Dis* 2014;58:164 .

⁵van den Bosch. *Clin Infect Dis* 2015;60(2):281.

Quality of Antimicrobial Prescribing^{1,2}

- Diagnostic criteria
- Empiric therapy
- Obtain cultures
- Culture-directed therapy
- Redundant regimen²
- IV/PO
- Document indication
- Duration of therapy

*

No consensus on what combination of these metrics constitutes “quality prescribing”

¹van den Bosch. *Clin Microbiol Infect* 2016; 888e1. ² Schultz. *Infect Control Hosp Epidemiol* 2014; 35:1229.

Test the utility of the quality indicator before widespread adoption^{1,2}

Applicability of generic quality indicators for appropriate antibiotic use in daily hospital practice: a cross-sectional point-prevalence multicenter study

C.M.A. van den Bosch ^{1,*}, M.E.J.L. Hulscher ², S. Natsch ³, J. Wille ⁴, J.M. Prins ¹, S.E. Geerlings ¹

4 /11 previously selected quality indicators were not clinically useful metrics

- Low applicability
- Low improvement potential
- Feasibility³

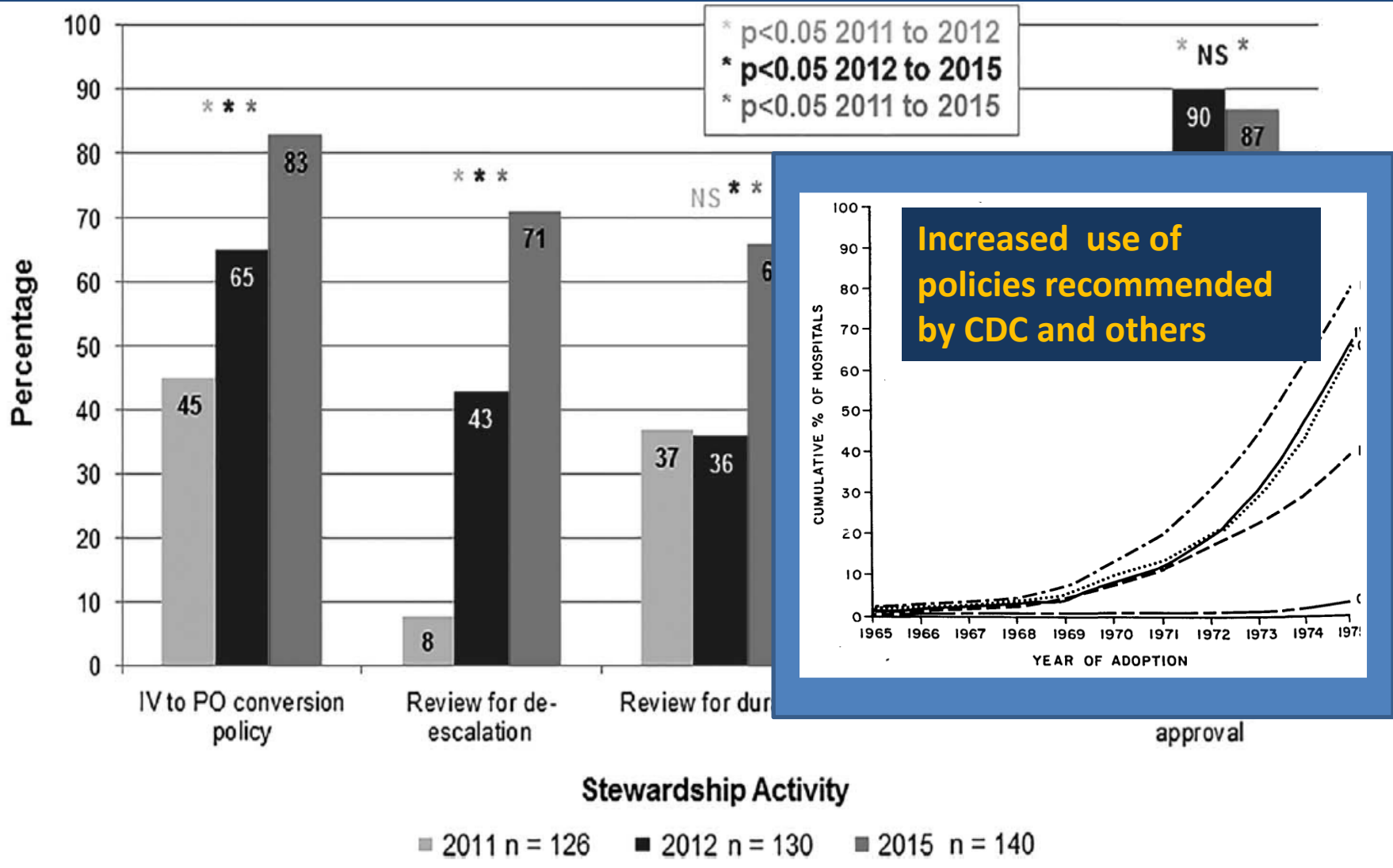
¹van den Bosch. *Clin Microbiol Infect* 2016; 22:888.e1. ²van den Bosch. *Clin Infect Dis* 2015;60(2):281. ³Moehring. *Clin Infect Dis* 2017; 64:377.

Which intervention (process measure)
should be selected?

Tailor interventions to the most
important issues at your site

Benchmarking Process Measures
Are there data?

Benchmarking Process Measures Veterans Health Administration (VHA)



“VHA is blazing a trail to improve patient safety through better antibiotic use”

... and working with the CDC to advance the science of using antibiotic use data to *guide action*

Quality Improvement

Defining key processes for improvement and metrics to monitor performance

Processes

- Interventions
- Quality indicators

**Demonstrated
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Outcomes

Outcomes^{1,2}

- Antibiotic use measures
- Patient outcomes
 - Mortality, LOS
 - Infection-related mortality
 - Unintended consequences (eg: *C. difficile*)
 - Not showing harm³
 - Conservable antibiotic days⁴
 - Unplanned readmission⁵
 - Unnecessary FQ days⁵
- Resistance
- Cost (value of healthcare)

¹Dodds Ashley. *Clin Infect Dis* 2014; (suppl 3):S112. ²Schuts. *Lancet Infect Dis* 2016; 16:847. ³Moehring. *Clin Infect Dis* 2017; 64:377. ⁴Morris. *Infect Control Hosp Epidemiol* 2012; 33:500. ⁵Werner. *BMC Infect Dis* 2011; 11:187.

Antibiotic use measures

National Healthcare Safety Network (NHSN)

- Antibiotic use (AU)
- Standardized Antimicrobial Administration Ratio (SAAR)

Application of NHSN data

Aggregate data

Helps an organization *focus resources*:

- an antibiotic or class of antibiotics
- that appear to be an outlier
- when compared with other facilities

*** *Your experiences with NHSN data?***

Antibiotic Use (AU) calculation

$$\text{AU} = \frac{\text{Days of therapy (DOT)}}{\text{Days present}^1 \text{ or \text{Admissions}^2}$$

¹Number of days patient spent any time in a specific unit or facility

²Number of patients admitted to an inpatient location in the facility

Standardized Antimicrobial Administration Ratio¹

Risk-adjusted

- Patient mix
- Hospital characteristics

SAAR Value =

$$\frac{\text{Observed number of abx days}}{\text{Predicted}^2 \text{ number of abx days}}$$

¹ Each SAAR is an observed to predicted ratio for a combination of antibiotics and patient care locations. SAAR (>1.0) indicates more AU than predicted; i.e., achieves statistical significance (different than 1.0). ²Statistically estimated from nationally aggregated data.

Application of SAARs

A Novel Metric to Monitor the Influence of Antimicrobial Stewardship Activities (SAAR)

Daniel J. Livorsi, MD, MSc;^{1,2} Erin O'Leary, MPH;³
Tamra Pierce, PharmD;⁴ Lindsey Reese, MD;⁴
Katharina L. van Santen, MSPH;³ Daniel A. Pollock, MD;³
Jonathan R. Edwards, MStat;³ Arjun Srinivasan, MD³

Prospective audit and feedback program:

- implementation of processes to **reduce broad-spectrum agents**
- greater involvement of an ID physician

.....associated with reduction in SAAR values across multiple antimicrobial categories

Application of SAARs

(Declines in broad-spectrum use, $p < .001$)

Findings consistent with efforts to:

- Encourage the prescription of more narrow-spectrum agents
- Not associated with increased LOS or mortality

Livorsi. *Infect Control Hosp Epidemiol* 2017; 38:721. Non-significant change between 2015-2014 of broad-spectrum antibiotics (BSA) for community infections, and all antibiotics.

Application of NHSN data (Aggregate data) (cont)

Does not:

- Inform about the appropriateness (quality) of antimicrobial prescribing

Reddy. *Expert Rev Anti Infect Ther* 2015; 13: 843.

Outcomes

Patient-specific outcomes

Metrics and quality improvement

Processes

**Demonstrated
linkage to
improve
outcome**



Outcomes
Patient-specific

Validity of linkage based on:

- scientific literature
- expert panel consensus

Current evidence on hospital antimicrobial stewardship objectives: a systematic review and meta-analysis

Schuts. *Lancet Infect Dis* 2016; 16:847.

- Clinical outcomes (mortality, LOS)
- Adverse events
- Bacterial resistance rates
- Costs

Literature
search

Quality indicators for
appropriateness of antibiotic use

Is there a
link?

Pre-determined outcomes:

- Clinical outcomes (mortality, LOS)
- Adverse events
- Bacterial resistance rates
- Costs

Literature search identified 14 quality indicators

Source	Comments	Number of Quality Indicators
<u>Expert consensus</u> – RAND-modified Delphi procedure ¹	Serial review of the literature & discussion, amongst experts	<u>Examples</u> <ul style="list-style-type: none"> • Empirical antibiotic therapy per local guide • De-escalation • ID Consultation • Others
<u>Expert consensus</u> – Antimicrobial stewardship consensus meetings ²		
Total		

¹ RAND: research and development corporation. van den Bosch. *Clin Infect Dis* 2015;60(2):281. ² Dellit. *Clin Infect Dis* 2007; 44:159 or stewardship guideline development group.

Next...Reviewed studies reporting the impact of the 14 quality indicators, on pre-defined outcomes*

Mortality, LOS
Adverse events
Bacterial resistance rates
Costs

Inclusion

- Hospital or long-term facilities, adults
- Randomized or non-RCTs controlled trials, until April 2014
- Interrupted time series
- Observational studies

Methods

- Pooled outcome data irrespective of study design or type of disorder
- Analyses to assess the risk for bias

Results

No published data identified of the impact of ASPs for the following.....*future study?*

Specific quality indicators (n=5)

1. Documented antibiotic plan
2. Blood cultures
3. Cultures from the site of infection
4. Local guide in agreement with national guidelines
5. Assessment of patients' adherence

Facility type

Stewardship objectives for long-term care facilities

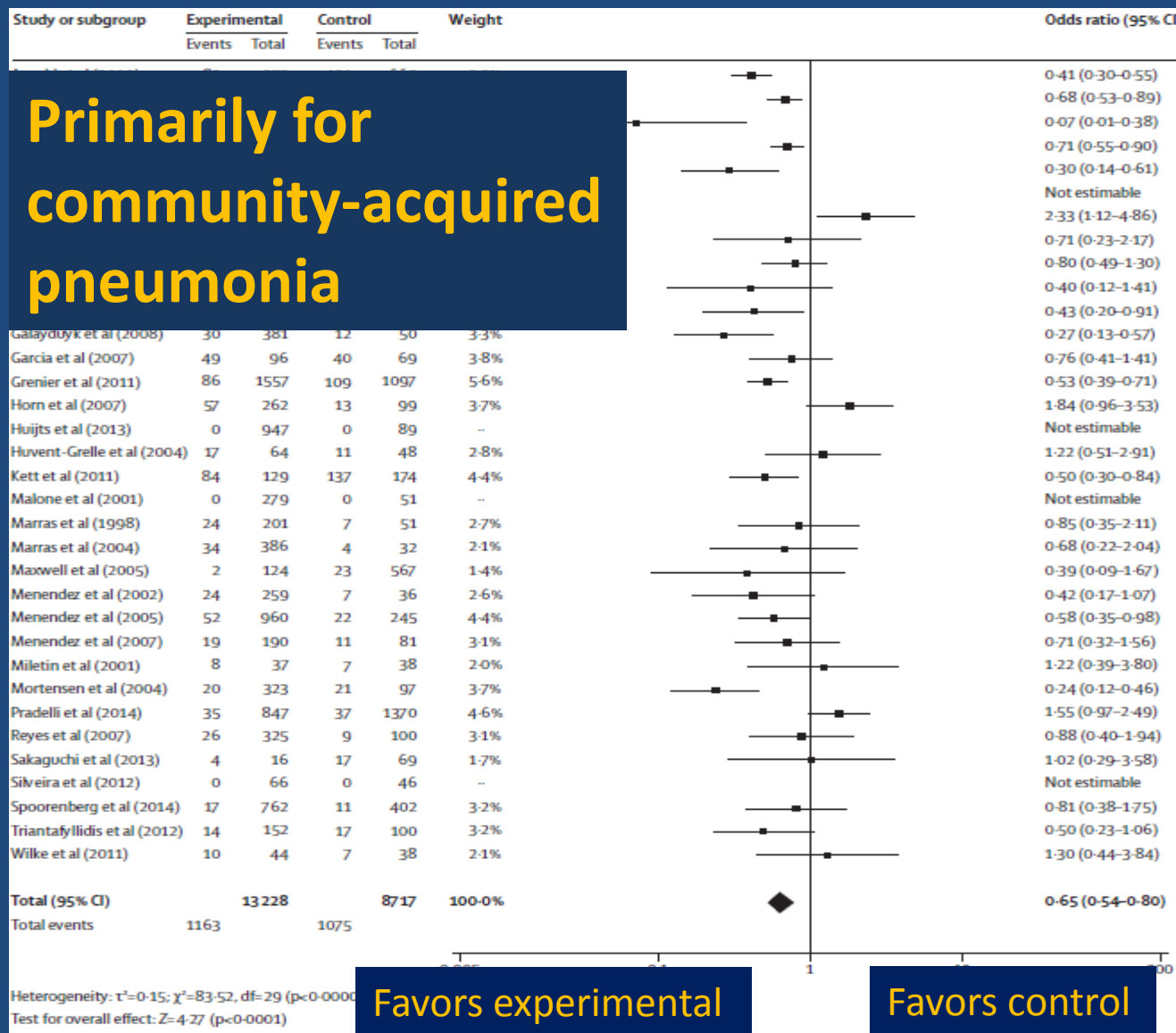
Remaining quality indicators: (n=9) with data evaluating the impact of ASPs

1. Empirical therapy according to the guidelines*
2. De-escalation of therapy*
3. Formal Bedside consultation by an ID specialist*

*Significant benefit demonstrated in >1 of 4 pre-defined outcomes; for ID bedside consultation, for *S. aureus bacteremia*. Schuts. *Lancet Infect Dis* 2016; 16:847.

Prescribing empirical antibiotic therapy according to guidelines (35% relative risk reduction for mortality¹)

Primarily for
community-acquired
pneumonia



¹ relative risk 0.65, 95% CI 0.54–0.80, $p<0.0001$. Schuts. *Lancet Infect Dis* 2016; 16:847.

De-escalation of therapy (56% relative risk reduction for mortality¹)

Study definition

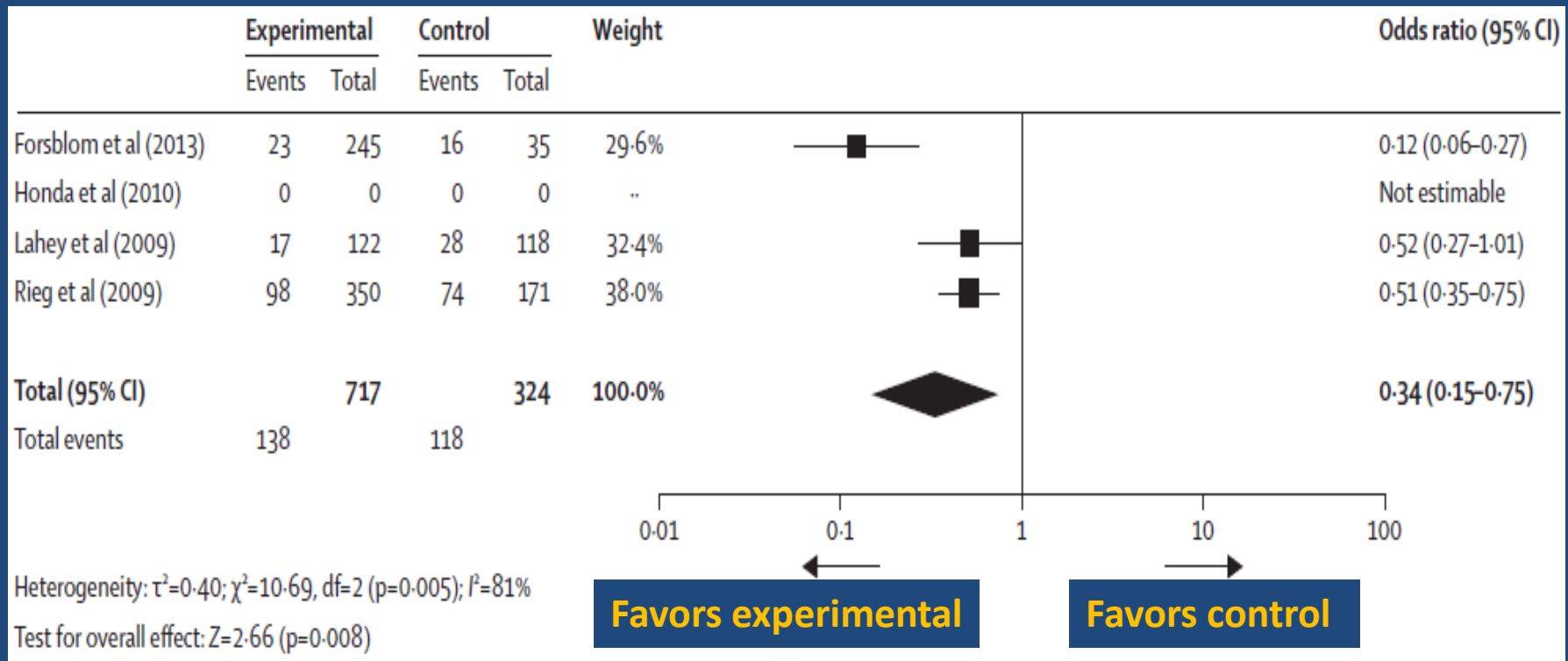
- Change to narrow-spectrum antibiotic or stop antibiotics as soon as culture results are available

Study design

- Mostly observational data

¹ RR (risk reduction) 0.44, 95% CI 0.30–0.66, $p < 0.0001$. Schuts. *Lancet Infect Dis* 2016; 16:847.

Bedside ID consultation (66% mortality reduction in *S. aureus* bacteremia)



Significant RRR of 66% (RR 0.34, 95% CI 0.15–0.75, $p=0.008$. Schuts. *Lancet Infect Dis* 2016; 16:847.

Overall, limited, low quality data

ASP's impact was positive for in >1 outcome

- Therapeutic drug monitoring
- Switch from IV to oral therapy
- Use of a list of restricted antibiotics

ASP's impact was less clear

- Adjusted of therapy per renal function
- Discontinuation of antibiotic therapy if infection is not confirmed
- Presence of a local guide

Limitations

- Interventions assessed separately
 - Interventions are generally bundled (in practice)
 - Combined effect of meeting several interventions could be greater than that of meeting one

Randomized multi-hospital trials needed:

Test the effectiveness of interventions on achieving meaningful stewardship outcomes

Comprehensive summary

- Interventions likely to be good starting points for any healthcare system:
 - *adherence to guidelines*
 - *de-escalation of therapy*
- Identified gaps in data linking interventions to practical outcomes

C. difficile infection

(*Lancet Infect Dis*; 2017)

Process

*Demonstrating
linkage to
outcome*

Outcomes

Effects of control interventions on *Clostridium difficile* infection in England: an observational study

Lancet Infect Dis 2017;
17: 411-21

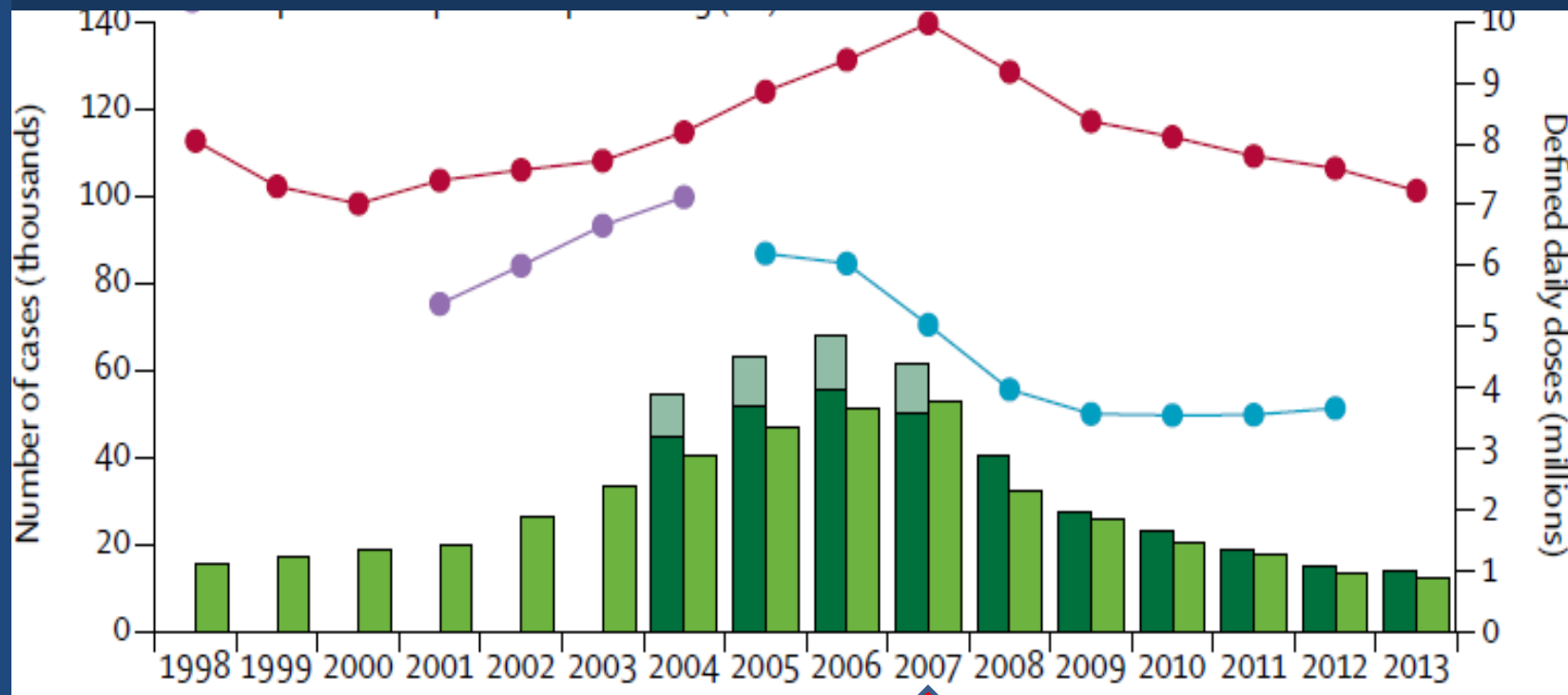
Conclusion:

....restriction of fluoroquinolone prescribing,
above other interventions

.....appears to explain the decline in incidence of
C. difficile infections

National recommendation (2007)

- Avoid clindamycin and cephalosporins
- Minimize use of fluoroquinolone, carbapenem and aminopenicillin
- Improved infection prevention and control activities



Dingle. *Lancet Infect Dis* 2017; 17: 411

Why did *C. difficile* infection decrease? (Two hypotheses)

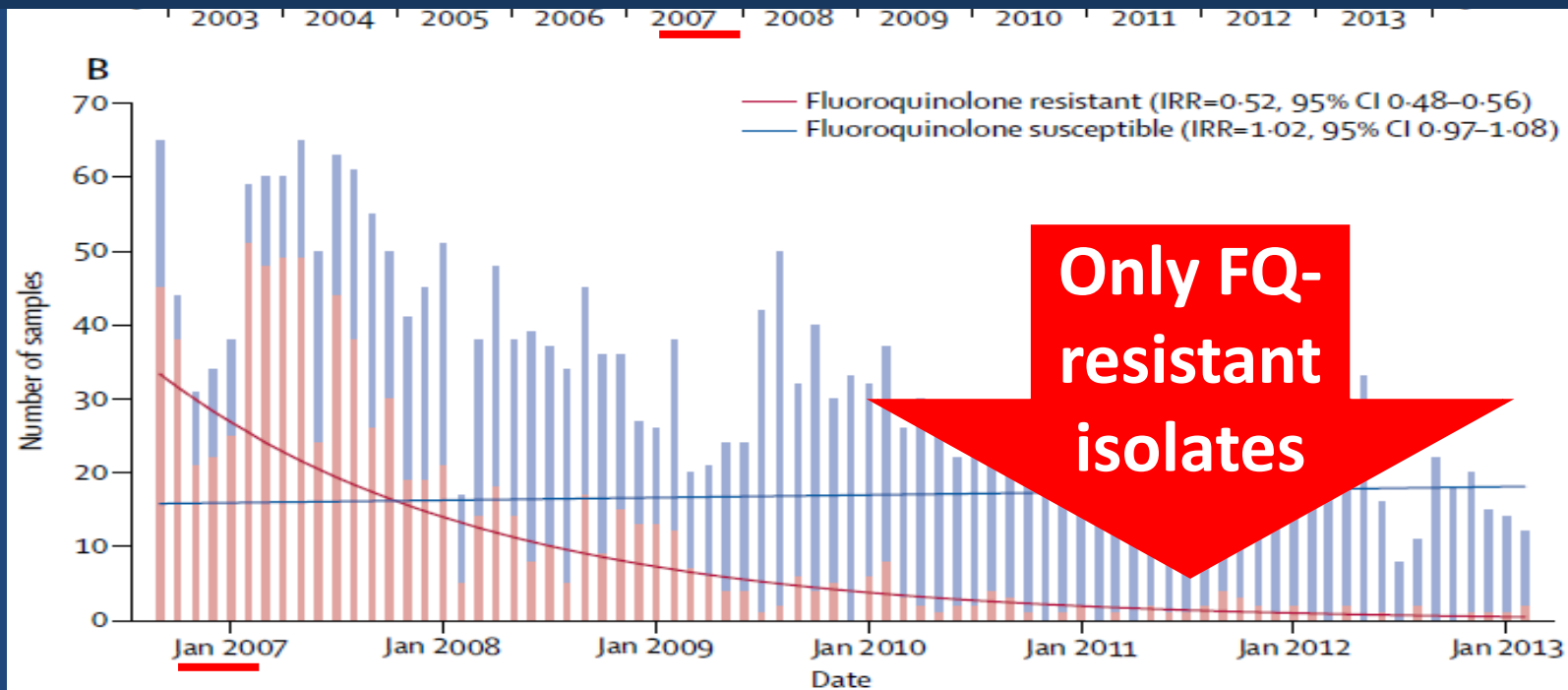
1. If declines were driven by reductions in use of particular antibiotics then:
then the incidence of *C difficile* infection caused by resistant isolates should decline faster than that caused by susceptible isolates across several genotypes.
2. If declines were driven by improvements in hospital infection control then:
transmitted (secondary) cases should decline regardless of susceptibility.

Methods /definitions

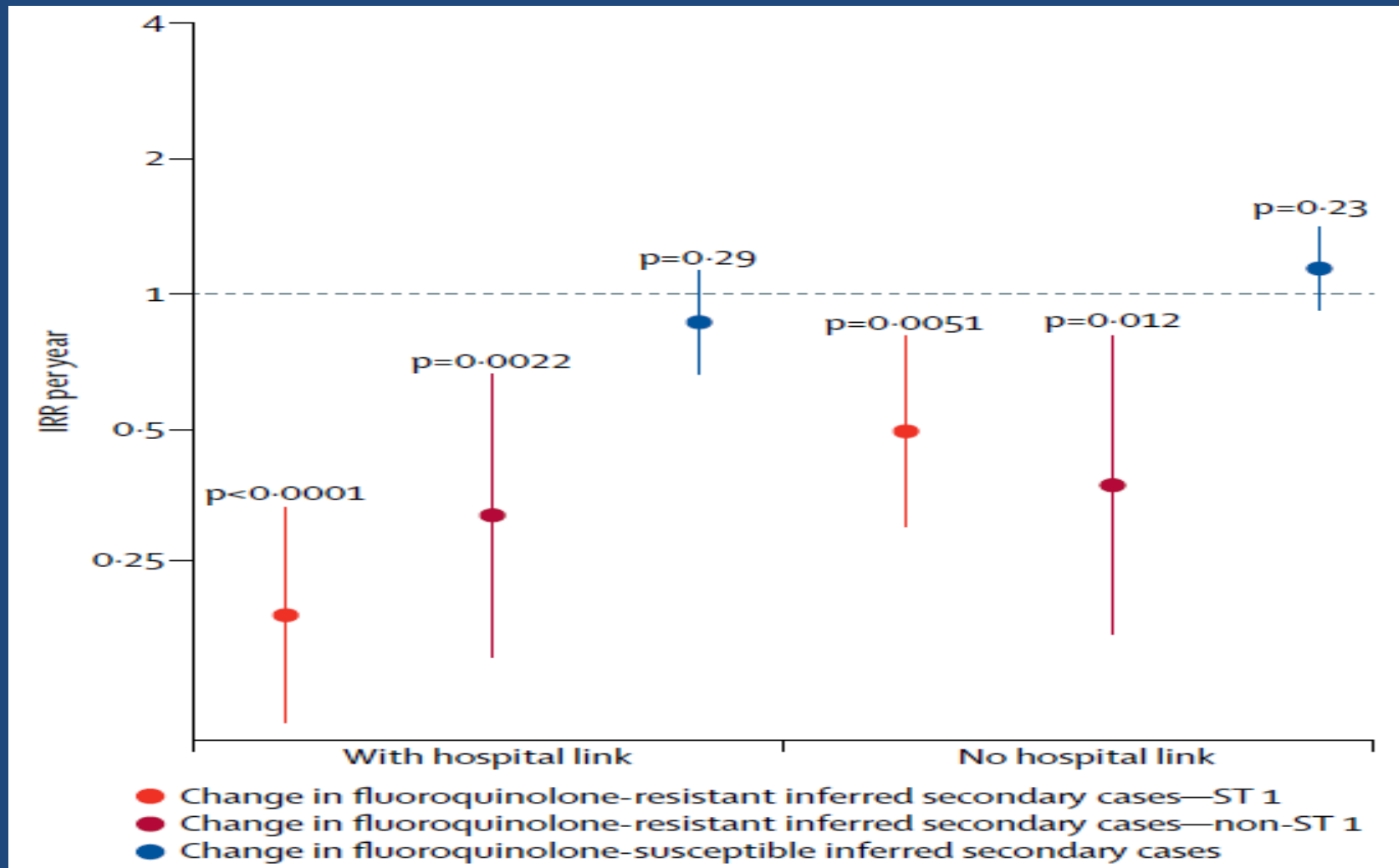
- Whole genome sequences:
 - clinical *C. difficile* isolates from symptomatic, unique patients during 2006-2013, (n=2021)
 - only sequence type, (n=261 isolates)
- Nosocomial transmission = subsequent infections from closely genetically related isolates

Whole genome sequence data suggests:

- FQ restriction plausibly played the most important part in the decline of *C. difficile* infection



Incidence of *C. difficile* infection only fell for secondary cases caused by FQ-resistant isolates



Limitations

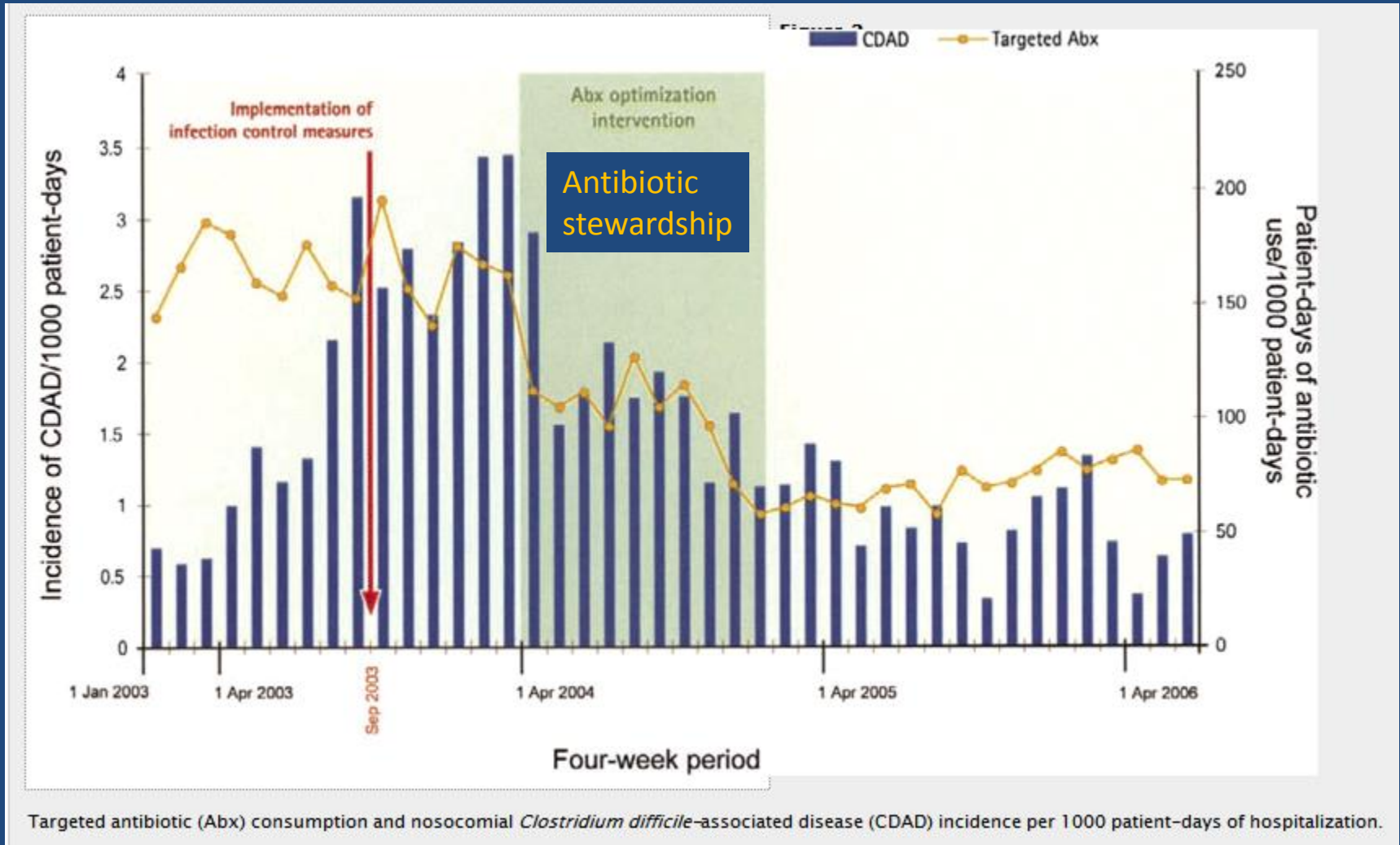
- Retrospective, quasi-experimental study

Nevertheless,

*these findings are
compelling, and consistent
with other data....*

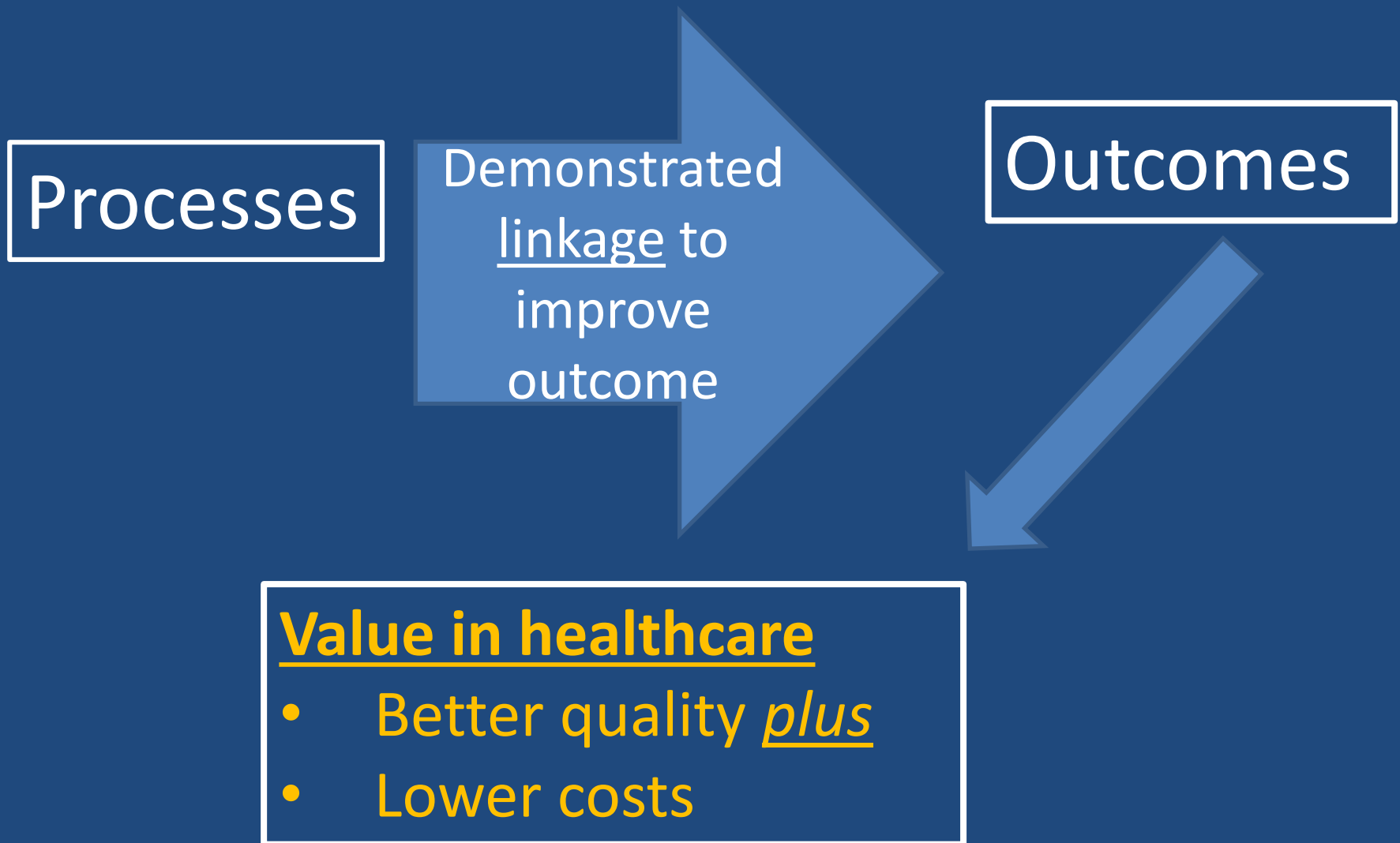
¹Donskey. *Lancet Infect Dis* 2017; 17:343. ² van Kleef. *Lancet Infect Dis* 2017; 17:478.

Antimicrobial Optimization Reduces *C. difficile* infection



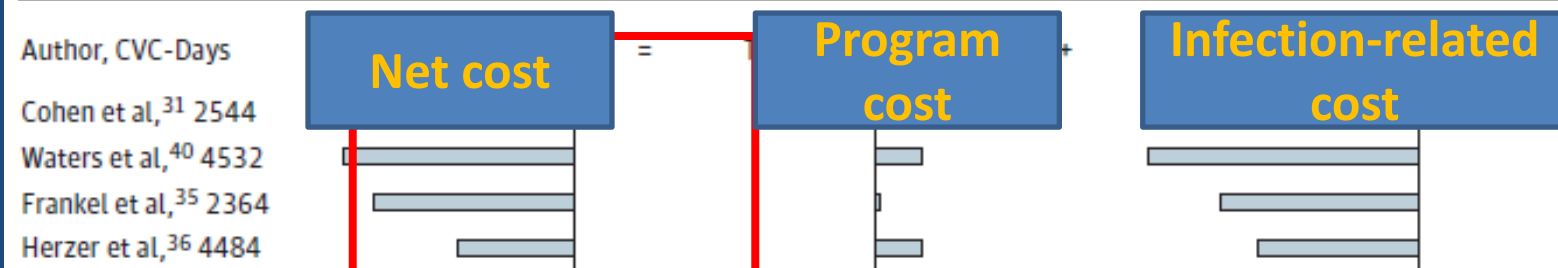
Valiquette L, et al. *Clin Infect Dis* 2007;45:S112-S121

Quality Improvement



Economic Evaluation of Quality Improvement Interventions for Bloodstream Infections Related to Central Catheters A Systematic Review

Figure 2. Net Costs Associated With Prevention of CLABSI and/or CRBSI Interventions From the Hospital Perspective Over 3 Years (2015 US Dollars)



- 57% fewer bloodstream infections
- Net savings in the millions....

...can be of high value

Demonstrating the Value of Antimicrobial Stewardship Programs to Hospital Administrators

- Reimbursement based on *quality* of care
- ASPs must:
 - expand beyond measures linked to cost and utilization

Regulatory of Quality Improvement Organizations

ASPs can demonstrate
value to administrators by:

Optimizing their hospital's
compliance with relevant
national quality indicators

*NHSN: National Healthcare Safety Network; CDC: Centers for Disease Control and Prevention. CMS: Centers for Medicare & Medicaid Services. AMA: American Medical Association; PCPI: Physician Consortium for Performance Improvement; (SC)AS: Laboratory Surgery Center. Nagel. *Clin Infect Dis* 2014;59(S3):S146.

Summary....*for now*....

- Aggregate NHSN antibiotic measures can be used by hospitals to:
 - identify antibiotics (or classes) that are outliers (compared to similar organizations)
 - does not inform on the quality of antibiotic use

Performance metrics

Identify important/problematic areas...

- WHAT are we trying to accomplish?
- WHY is it important?
- WHO is the specific target population?
- WHAT are our measureable goals?
- HOW will we know our changes are working?
- HOW do we know if the intervention is harmful?

- HOW completely was intervention implemented?
- VALIDITY of data - alternate reasons for results

Finally, more studies....

- Link between ASP interventions (processes) and improved patient outcomes
- Value of ASPs
 - Health outcomes achieved per dollar spent

11 years after infection control programs were mandated by the Joint Commission.....

Editorial

INFECTION CONTROL 1987/Vol. 8, No. 6

Peter C. Fuchs, MD, PhD

Will the Real Infection Rate Please Stand?

Acknowledgements

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¹ John H. Stroger Jr. Hospital of Cook County

Questions / comments?

Your experiences with:

Joint Commission Survey?

NHSN usage measures?