

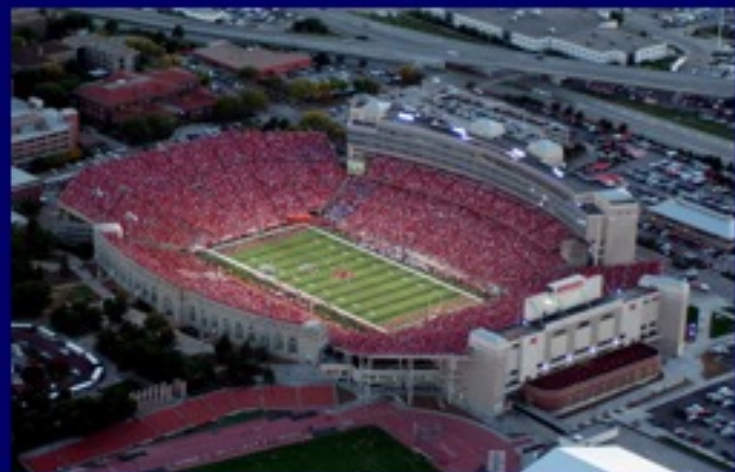
Technologic Innovations to Prevent Catheter-Related Bloodstream Infection

Mark E. Rupp, MD

Professor & Chief, Infectious Diseases
Director, Infection Control & Epidemiology
University of Nebraska Medical Center

Hosted by Bruce Gamage
Provincial Infection Control Network of British Columbia

Nebraska



Clinical Significance of CLA-BSI

- 78,000 central line-associated bloodstream infections (CLA-BSI) are estimated to occur yearly in United States hospitals and dialysis units.
- 2013 NHSN report from 4,567 US facilities, mean CLA-BSI rate in critical care units ranged from 0.0 – 3.0/1000 CVC d.
- CLA-BSI are associated with an estimated mortality rate of 12.3% and excess healthcare costs between \$7,288 and \$29,156 per episode.



International Nosocomial Infection Control Consortium (INICC) report, data summary of 43 countries for 2007-2012. Device-associated module



Rosenthal VD, et al. AJIC 2014

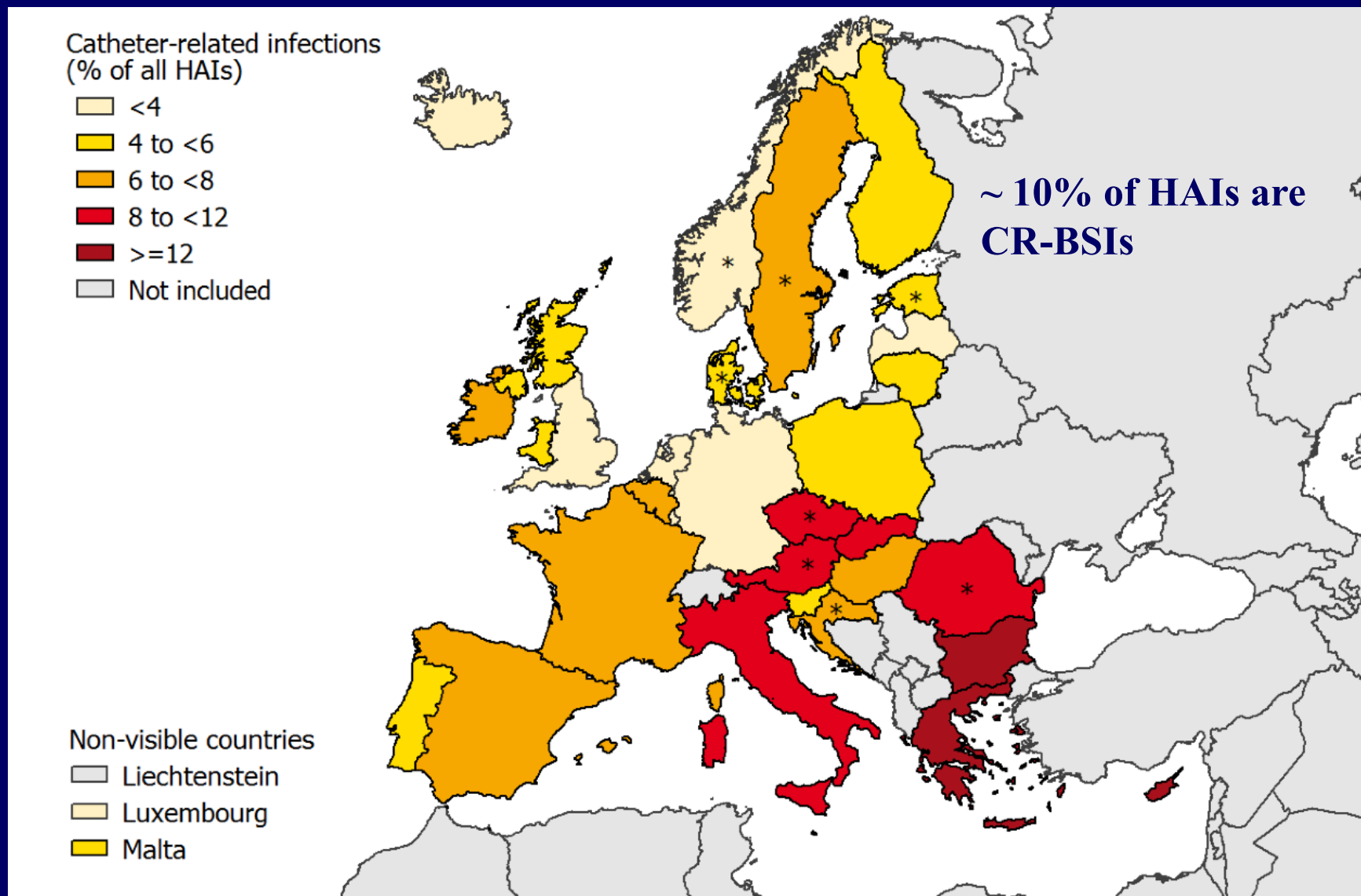
- **Countries Represented:** Argentina, Bolivia, Brazil, Bulgaria, China, Colombia, Costa Rica, Cuba, Cyprus, Dominican Republic, Ecuador, Egypt, El Salvador, Greece, Honduras, India, Iran, Saudi Arabia, Kosovo, Kuwait, Lebanon, Lithuania, Macedonia, Malaysia, Mexico, Mongolia, Morocco, Pakistan, Panama, Peru, Philippines, Poland, Puerto Rico, Romania, Russia, Serbia, Slovakia, Sudan, Thailand, Tunisia, Turkey, United Arab Emirates, Uruguay, Venezuela, Vietnam

Comparison of device-associated health care–associated infection rates per 1,000 device days in the ICUs of the INICC (2007-2012) and the U.S. NHSN (2012)

DA-HAI per Type of ICU	INICC 2007-2012 pooled mean (95% CI)	U.S. NHSN 2012 pooled mean (95% CI)
Medical cardiac ICU		
CLABSI	3.5 (3.1-3.9)	1.1 (1.0-1.1)
CAUTI	5.9 (5.4-6.4)	2.2 (2.0-2.3)
VAP	11.5 (10.5-12.5)	1.0 (0.8-1.1)
Medical and surgical ICU		
CLABSI	4.9 (4.8-5.1)	0.9 (0.9-1.0)
CAUTI	5.2 (5.2-5.8)	1.2 (1.2-1.3)
VAP	16.5 (16.1-16.8)	1.1 (1.0-1.2)
Pediatric ICU		
CLABSI	6.1 (5.7-6.5)	1.4 (1.3-1.6)
CAUTI	5.6 (5.1-6.1)	2.7 (2.5-3.0)
VAP	7.9 (7.4-8.4)	0.8 (0.6-0.9)
Newborn ICU (1,501-2,500 g)		
CLABSI	4.8 (3.7-6.1)	0.6 (0.5-0.8)
VAP	10.7 (8.4-13.4)	0.2 (0.1-0.5)

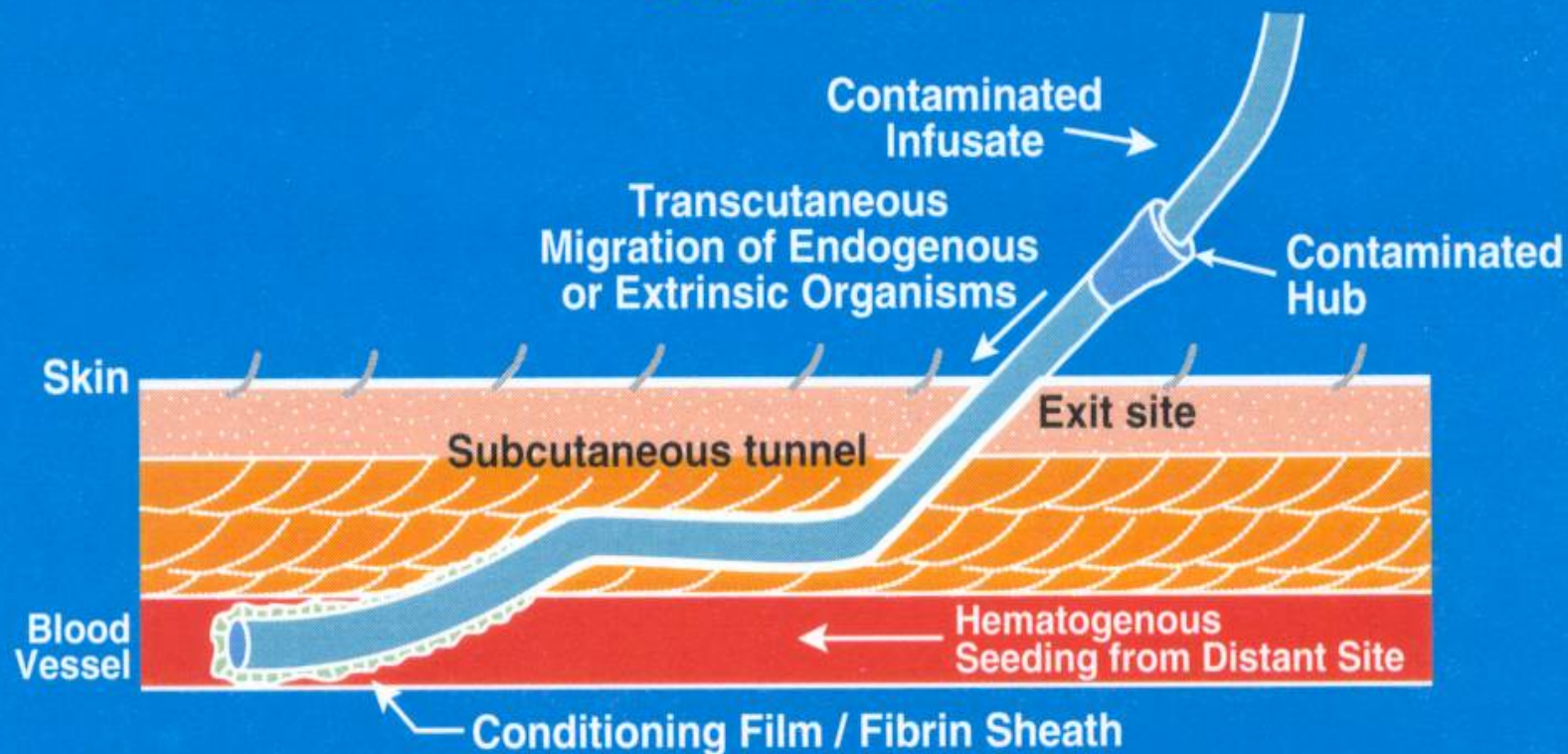
CAUTI, catheter-associated urinary tract infection; CI, confidence interval; CLABSI, central line-associated bloodstream infection; DA-HAI, device-associated health-care-associated infection; ICU, intensive care unit; INICC, International Nosocomial Infection Control Consortium; NHSN, National Healthcare Safety Network; VAP, ventilator-associated pneumonia.

Relative frequency of CR-BSI as a total of all HAIs by country (ECDC PPS 2011-2012)

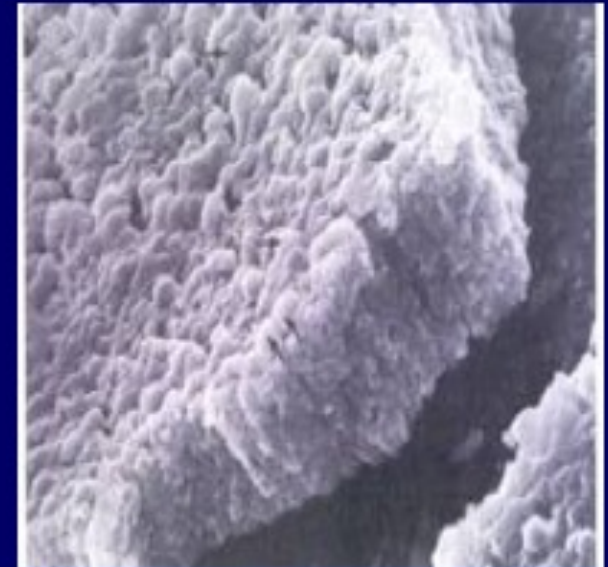
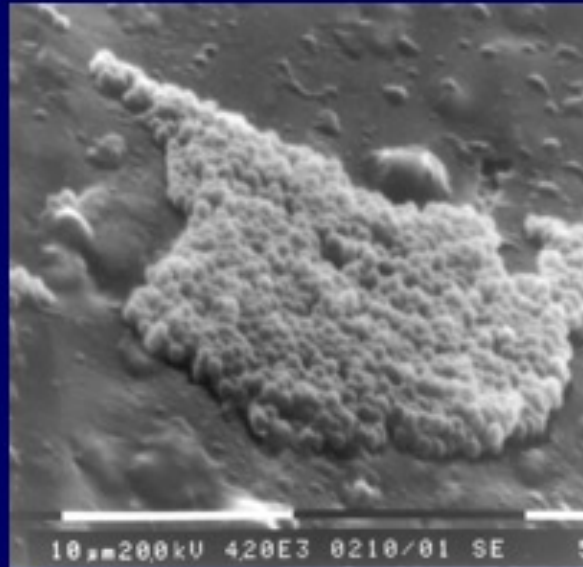


Pathogenesis of CVC-Associated BSI

Potential Sources for Bacterial Contamination of Intravascular Catheters



Pathogenesis of CVC-Associated BSI



Mature biofilm-associated infection with diverse population of cells including “persistor cells” is very difficult to eradicate with catheter in place

Prevention of CR-BSI

Practice Associated Interventions



Practice Associated Interventions

- Education & Training
- Staffing Levels
- Insertion Procedures
 - Full Sterile Barriers & Checklist
- Post Insertion Care
 - Dressing Integrity
 - Aseptic Access Technique (scrub the hub)
 - Discontinuing unneeded catheters



The NEW ENGLAND JOURNAL of MEDICINE

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DECEMBER 28, 2006

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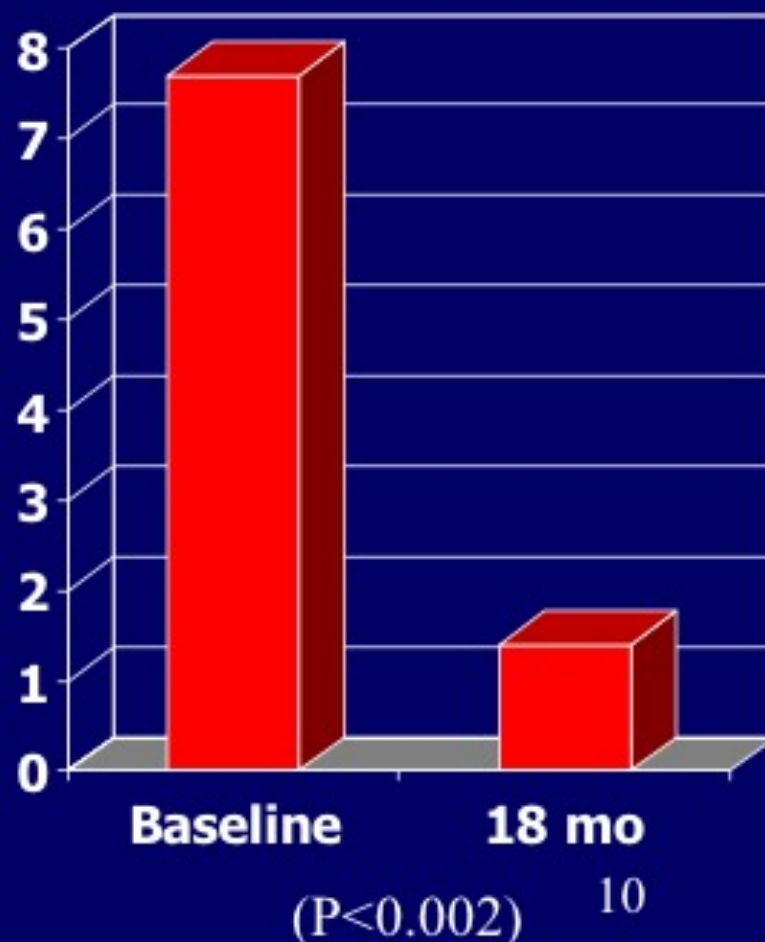
An Intervention to Decrease Catheter-Related Bloodstream Infections in the ICU

Peter Pronovost, M.D., Ph.D., Dale Needham, M.D., Ph.D., Sean Berenholtz, M.D., David Sinopoli, M.P.H., M.B.A., Haitao Chu, M.D., Ph.D., Sara Cosgrove, M.D., Bryan Sexton, Ph.D., Robert Hyzy, M.D., Robert Welsh, M.D., Gary Roth, M.D., Joseph Bander, M.D., John Kepros, M.D., and Christine Goeschel, R.N., M.P.A.

Intervention in 108 ICUs:

- Daily Goals Sheet
- Hand Hygiene
- Full Sterile Barrier Precautions
- Chlorhexidine Antiseptic
- Avoidance of the Femoral Site
- Removal of CVCs asap

Mean BSI/1000 CVC d



Nebraska Medical Center CVC Insertion Kit



The Nebraska Medical Center, 987400 Nebraska Medical Ctr., Omaha NE 68198-7400 Ph: 402-559-4...

Patient Information

Patient Name	Sex	DOB
[REDACTED]	Female	6/25/1996

CVC Checklist Note signed 1 **RN at 6/12/2014 6:28 PM**

Author:	Author	Registered Nurse
RN	Type:	
Filed: 6/12/2014 6:28 PM	Note Time: 6/12/2014 6:27 PM	Co-signer: [REDACTED], RN at 6/12/2014 6:38 PM

Central Venous Catheter Checklist*

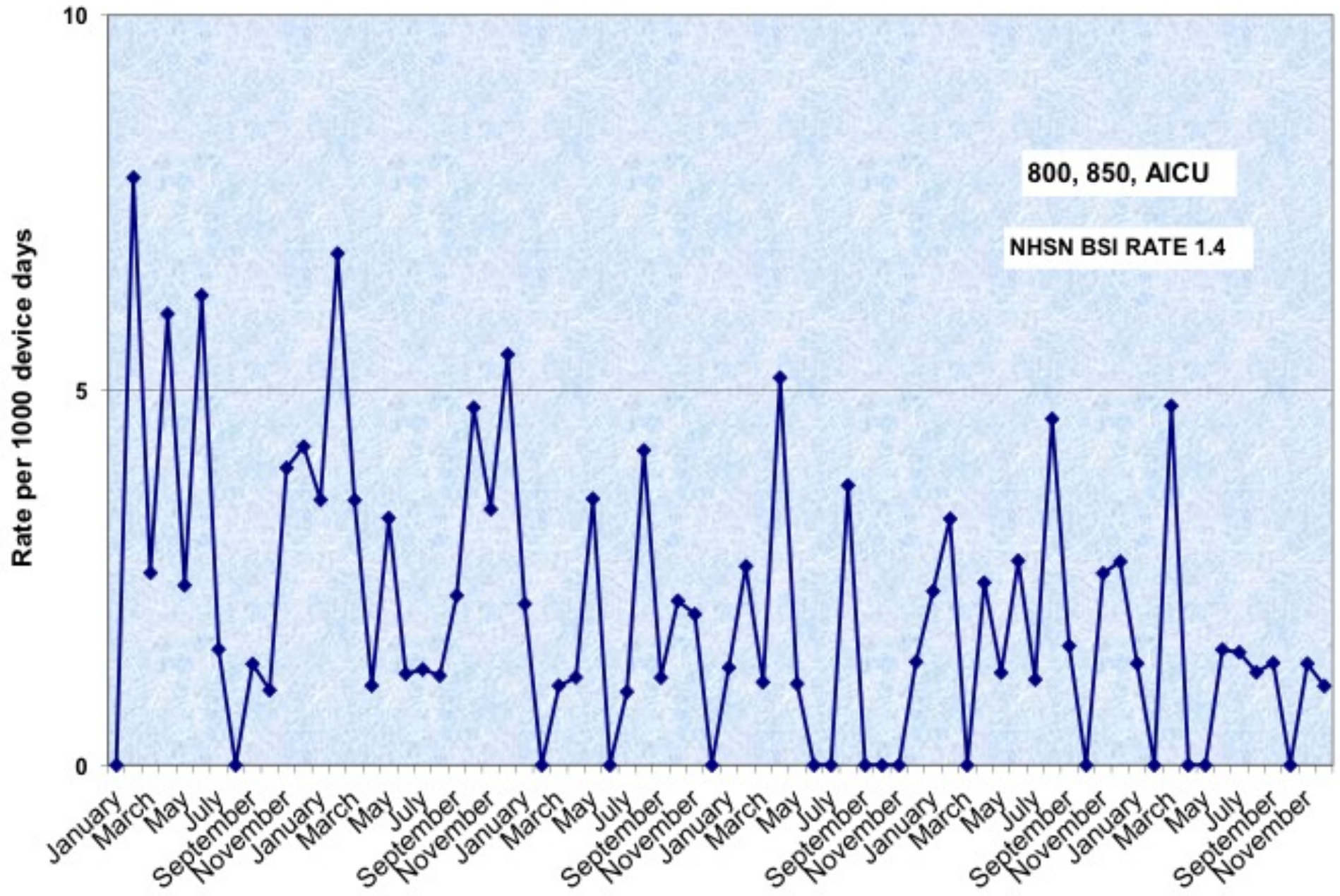
10000 [REDACTED] 10000 [REDACTED]
Date of Birth: 6/25/1996 Gender: female

Unit / Room: 800 ICU
Type of line inserted: PCC insertion
Is the line tunneled or un-tunneled or port?
Reason for insertion: New indication
If suspected central-line associated infection, was the central line exchanged over a guidewire? N/A
Name of person performing procedure: S. Rouse RN
Occupation of inserter: PICC Team
Supervising Physician / Provider
Inserter performed hand hygiene prior to line insertion: Yes
All personnel in room must wear cap, mask & gloves: Yes
Operator(s) don cap, mask & eye protection: Yes
Skin preparation: Chlorhexidine gluconate
If Chlorhexidine was not used, was there a contraindication for Chlorhexidine? Yes
Was skin prep dry at time of first puncture? (if not directly observed, ask inserter) Yes
Operator(s) don sterile gown & sterile gloves? Yes
Drape site with full-body sterile drapes (head to toe): Yes
Sterile field maintained throughout procedure: Yes
Did the insertion attempt result in a successful central line placement? No
Dressing applied: Yes
Observer was present throughout the procedure? Yes
Comments (pertaining circumstances, breaks in protocol, rationale for review):

Observer Signature: [REDACTED], RN 6:27 PM 6/12/2014

*Responses in Bold/Red are required elements of the CLABSI prevention bundle

**Combined Adult ICU Blood Stream Infection Rates
January 2007 through December 2012**



What if Practice Measures Aren't Working??????



Prevention of CR-BSI

Technologic Innovations



ARROWgard Blue PLUS[®]
Central Venous Catheters



Behavioral Change vs. Technology

“If you can choose between education and influencing human behavior or introduction of a gizmo, choose the gizmo everytime.”

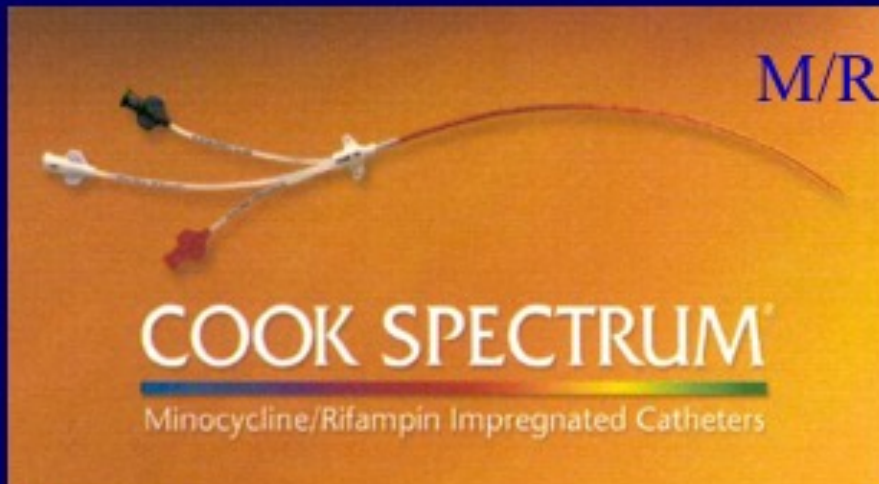
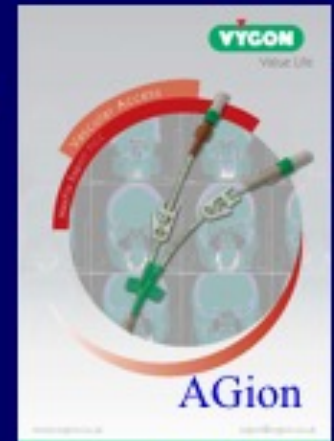
Bob Weinstein

Commercially Available Antimicrobial Central Venous Catheters



CHG/SS

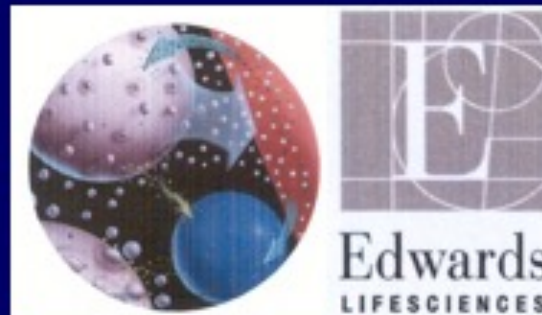
ARROW⁺gard Blue PLUS[®]
Central Venous Catheters



M/R

COOK SPECTRUM[®]

Minocycline/Rifampin Impregnated Catheters



Silver/Platinum/
Carbon (Silver
Iontophoretic; Vantex)

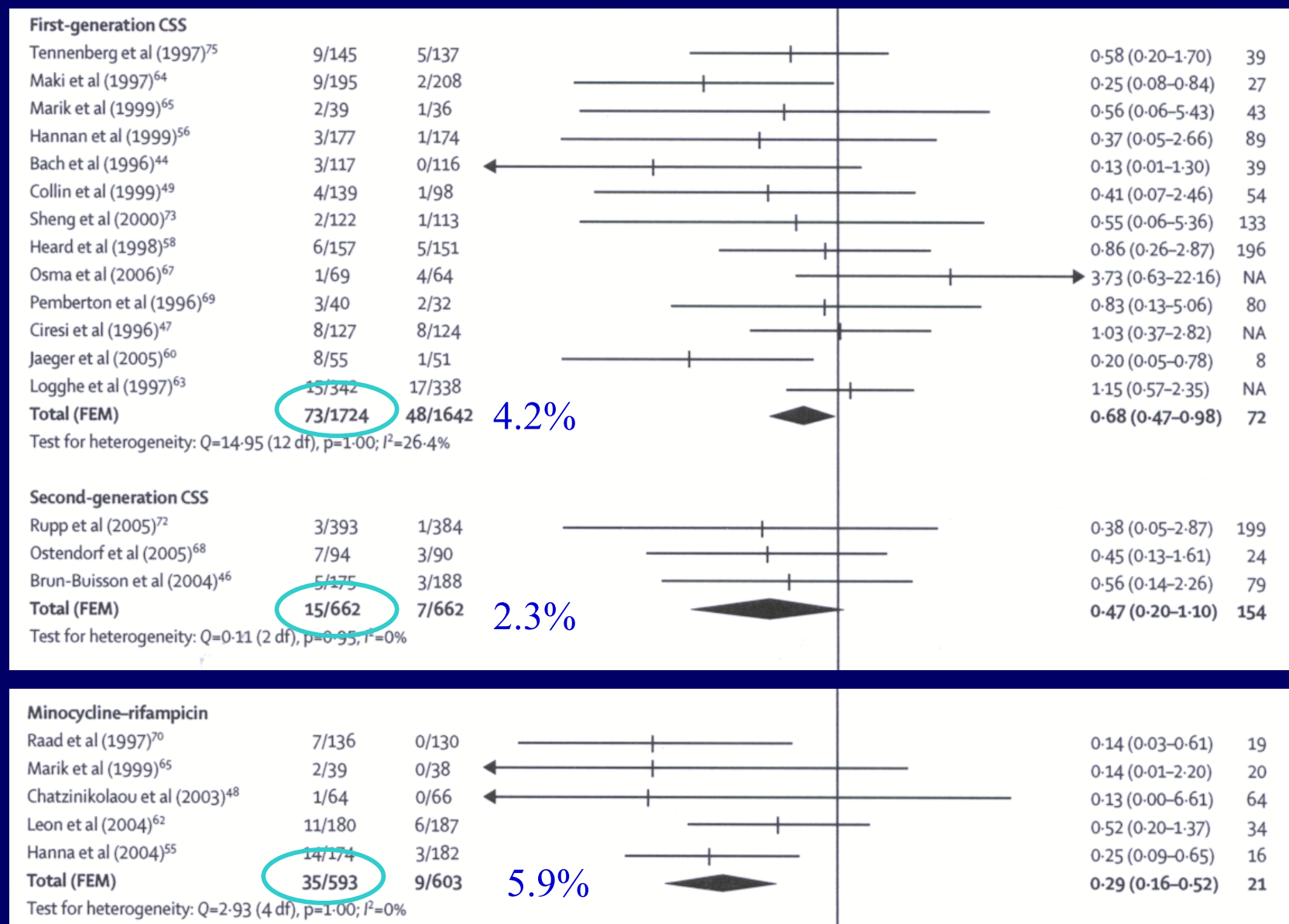


Hydrocath Assure (BD)
Benzalkonium
AMC Thromboshield (Edwards)
Benzalkonium Heparin



Multistar
Miconazole/Rifampin
(Vygon)

Do antimicrobial-coated catheters prevent BSI?



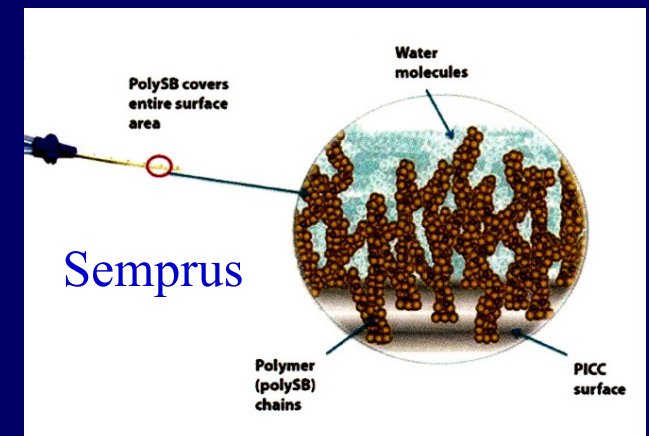
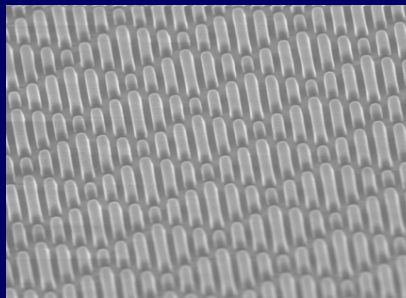
0.68

0.47

0.29

Novel Antimicrobial Coatings & CVCs

- 5-Fluorouracil
- Rifampin-Miconazole
- Silver Nanoparticles
- Chlorhexidine/Minocycline/Rifampin
- Gentian violet/Chlorhexidine
- Surface Pattern (Sharklet)
- Polymeric sulfobetaine (polySB)
- Heated CVCs



Preservation of Dressing Integrity

- Dressing disruption is a major risk factor for catheter-related infections

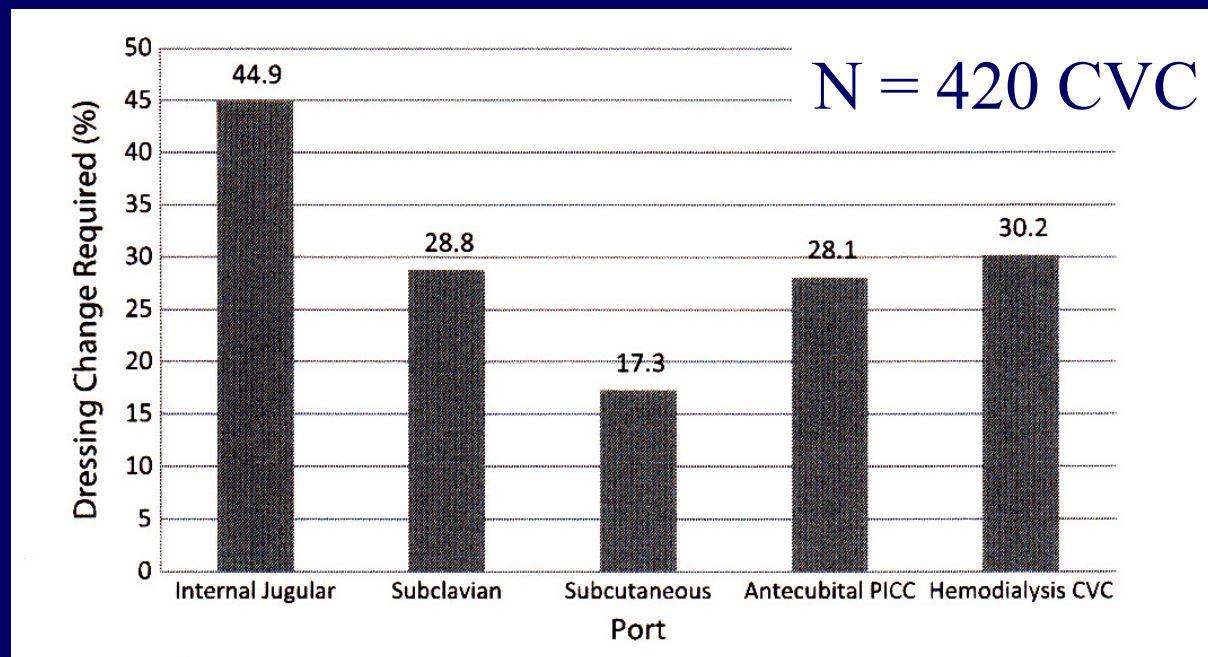
Table 5. Association between dressing disruption and catheter colonization or infection (unadjusted and adjusted marginal Cox model)

	Catheter Colonization $\geq 10^3$ colony-forming units/mL		Catheter-Related Bloodstream Infection		Major Catheter-Related Infection	
	HR (95% CI)	p	HR (95% CI)	p	HR (95% CI)	p
Unadjusted						
First	1.64 (1.13–2.39)	.01	3.15 (0.67–14.79)	.15	2.66 (0.50–14.26)	.25
Second disruption	1.52 (1.14–2.04)	.005	5.18 (1.85–14.48)	.002	4.31 (1.39–13.41)	.012
Final disruption	13.54 (10.17–18.04)	<.0001	14.90 (6.40–34.64)	<.0001	13.41 (5.17–34.75)	<.0001
Adjusted*						
First disruption	1.30 (0.90–1.87)	.16	2.65 (0.67–10.56)	.17	1.94 (0.50–7.48)	.33
Second disruption	1.16 (0.87–1.55)	.33	4.49 (1.71–11.79)	.002	3.26 (1.18–9.02)	.023
Final disruption ^b	13.99 (9.88–19.82)	<.0001	18.11 (5.66–57.88)	<.0001	12.51 (3.95–39.62)	<.0001

- The number of dressing disruptions was related to increased risk of colonization and bloodstream infection ($P < 0.001$)

Hospital-wide assessment of compliance with central venous catheter dressing recommendations

Mark E. Rupp MD^{a,b,*}, Kyle Cassling BA^a, Hayley Faber BS^a, Elizabeth Lyden MS^c, Kate Tyner RN^b, Nedra Marion RN^b, Trevor Van Schooneveld MD^{a,b}

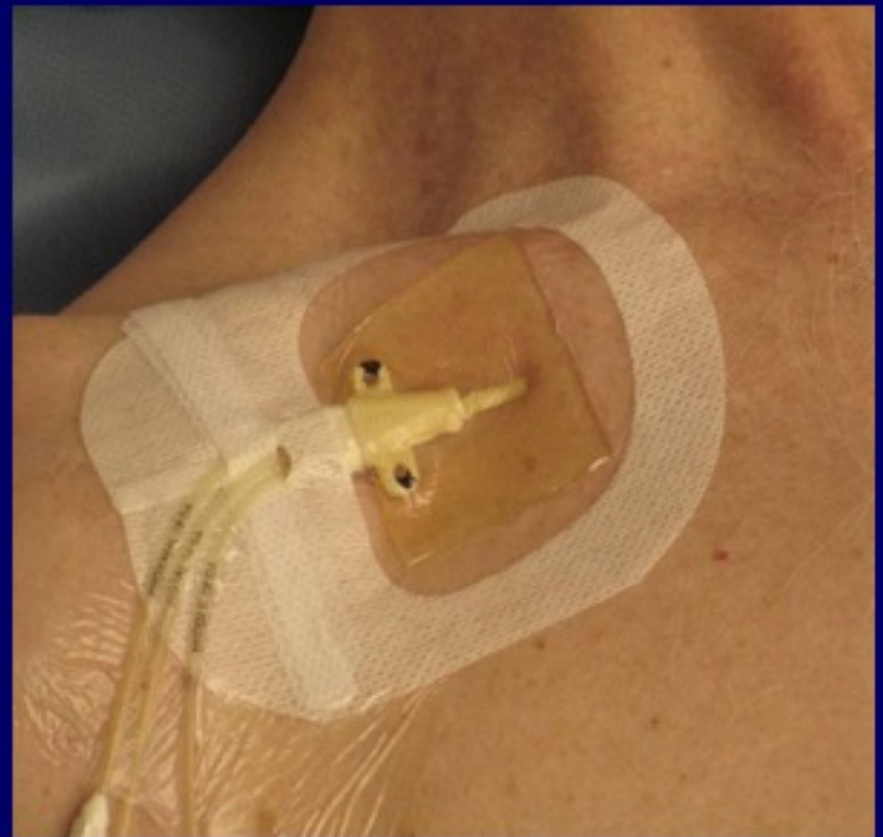


- On any given day approximately 31% of dressings were suboptimal and in need of change
- Reasons: 69% blood under dressing, 25.4% edge lift, 5.4% moisture under dressing

Chlorhexidine Impregnated CVC Dressings



“Biopatch”

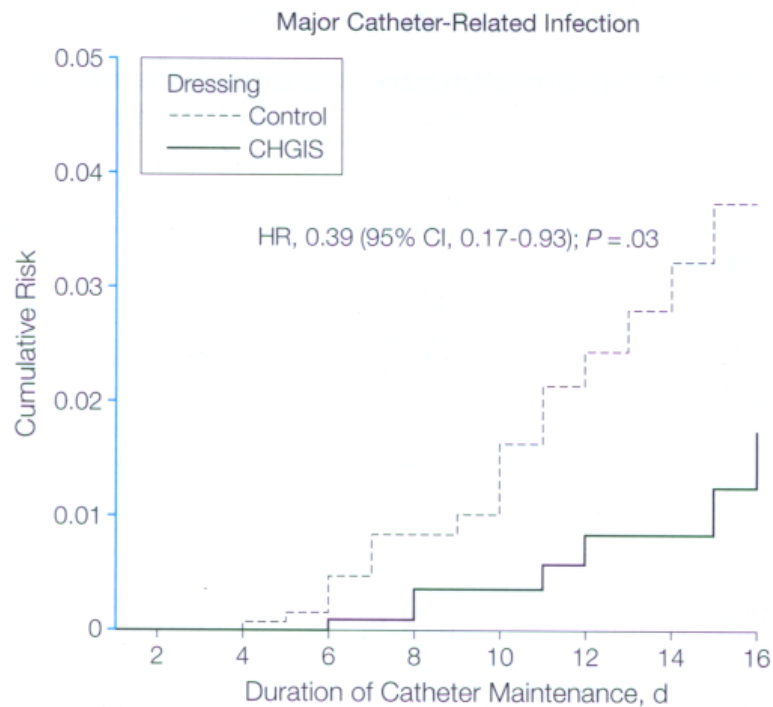


“Tegaderm CHG”

Chlorhexidine-Impregnated Sponges and Less Frequent Dressing Changes for Prevention of Catheter-Related Infections in Critically Ill Adults

A Randomized Controlled Trial

Timsit, et al. JAMA, 2009

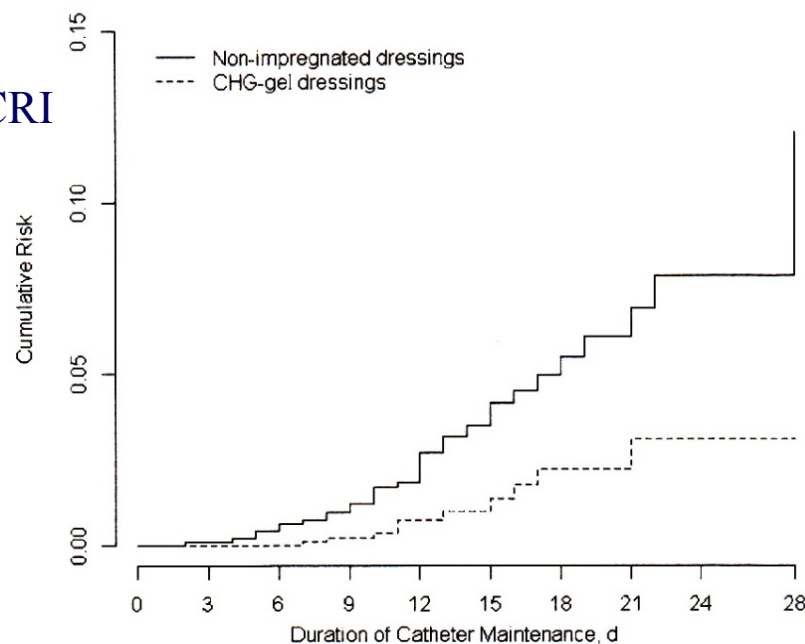


- CR-BSI: 1.4/1000 CVC d vs 0.4/1000 CVC d ($P < 0.005$)
- No significant difference between 3d and 7d dressing changes
- Full sterile barrier precautions used
- Site prep with 4% povidone-iodine soln & PI/Etoh

Randomized Controlled Trial of Chlorhexidine Dressing and Highly Adhesive Dressing for Preventing Catheter-related Infections in Critically Ill Adults

Timsit, et al. Crit Care Med, 2013

Major CRI



- CR-BSI: 1.3/1000 CVC d vs 0.5/1000 CVC d (P= 0.02)
- Major-CRI: 2.1/1000 CVC d vs 0.7/1000 CVC d (P=0.0006)
- Highly adhesive dressings decreased dressing detachment rate (71.9% vs 64.3%; P<0.0001) but increased rate of colonization HR 1.65, 95%CI 1.21-2.26, P=0.0016)

Scrub the Hub!

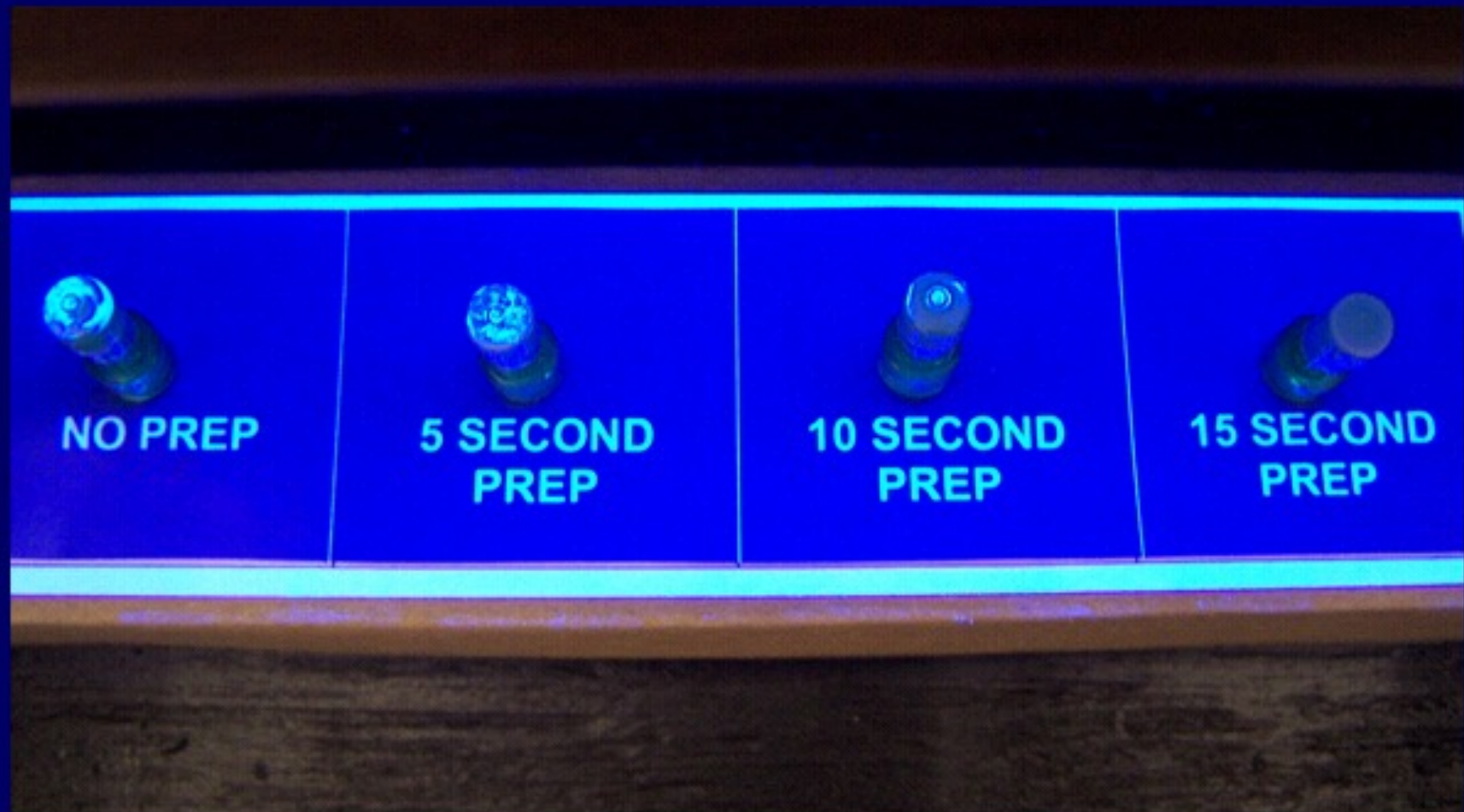
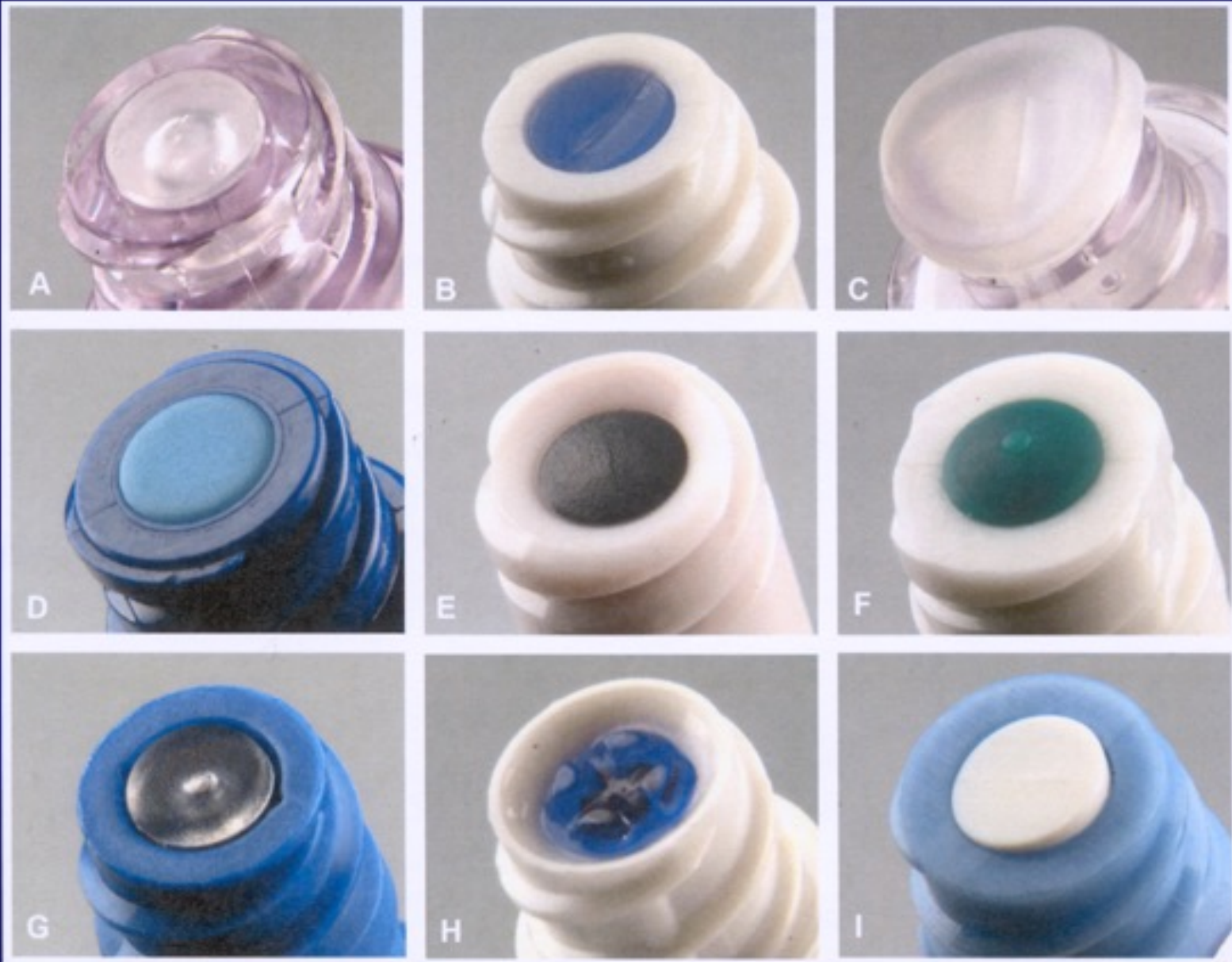


Figure: Courtesy Kristina Bryant, Kosair Children's Hosp

Not All Mechanical Valves are Created Equal



Needleless Connector Valves Linked to Increased CLA-BSI



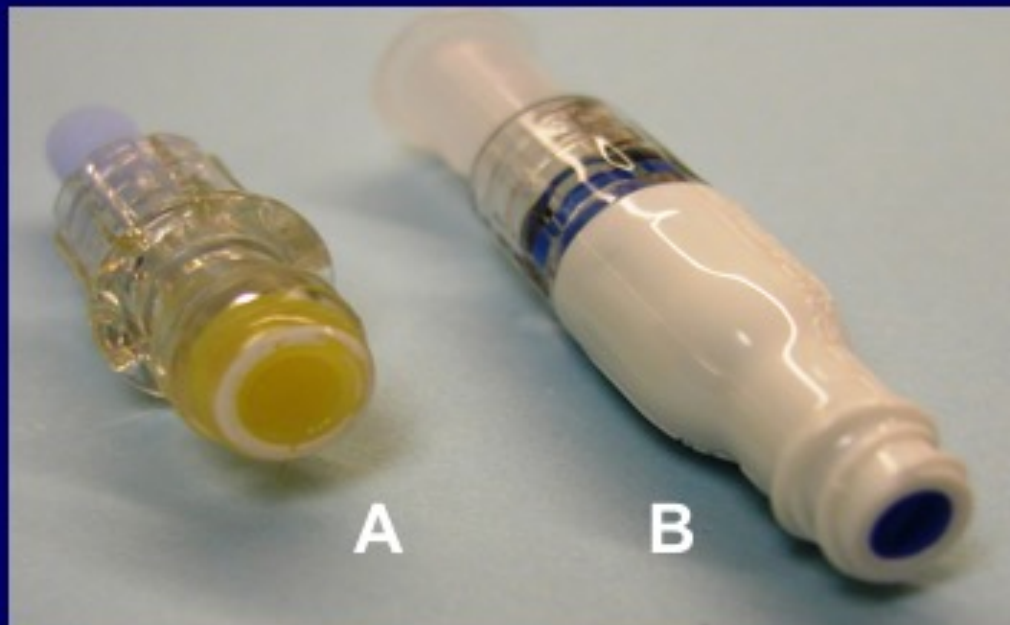
- Maragakis: ICHE, 2006
- Rupp: Clin Infect Dis, 2007
- Salgado: ICHE, 2007
- Field: ICHE, 2007
- Toscano: AJIC, 2009
- Jarvis: Clin Infect Dis, 2009

(Figure from Jarvis, Infect Control Today, 2010)

Outbreak of Bloodstream Infection Temporally Associated with the Use of an Intravascular Needleless Valve

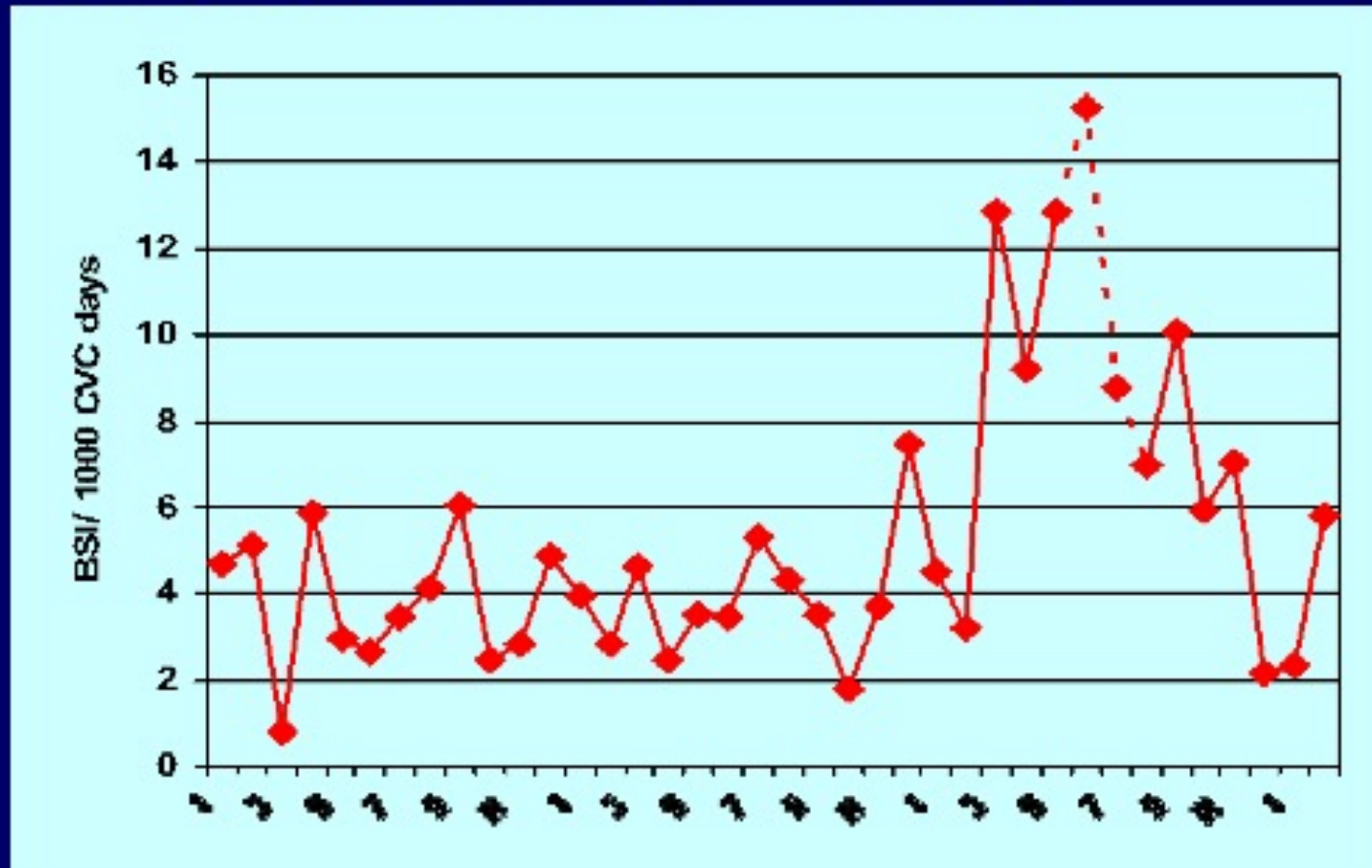
Mark E. Rupp, Lee A. Sholtz, Dawn R. Jourdan, Nedra D. Marion, Laura K. Tyner, Paul D. Fey, Peter C. Iwen and James R. Anderson

Clin Infect Dis 2007



A: Interlink IV Access System, Baxter
B: SmartSite Plus, Alaris Medical Systems

BSI Critical Care Units



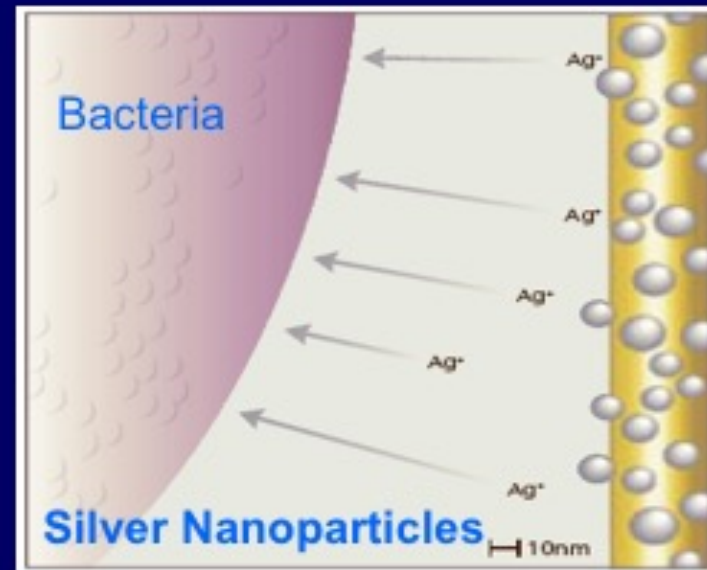
8 critical care areas, 132 beds.

Baseline: 38,250 CVC days, rate 3.87/1000 CVC d

Outbreak: 10,340 CVC d, rate 10.64/1000 CVC d
(2.82 fold increase) ($p < 0.0001$)

Post removal: rate 5.59/1000 CVC d ($p = 0.02$)

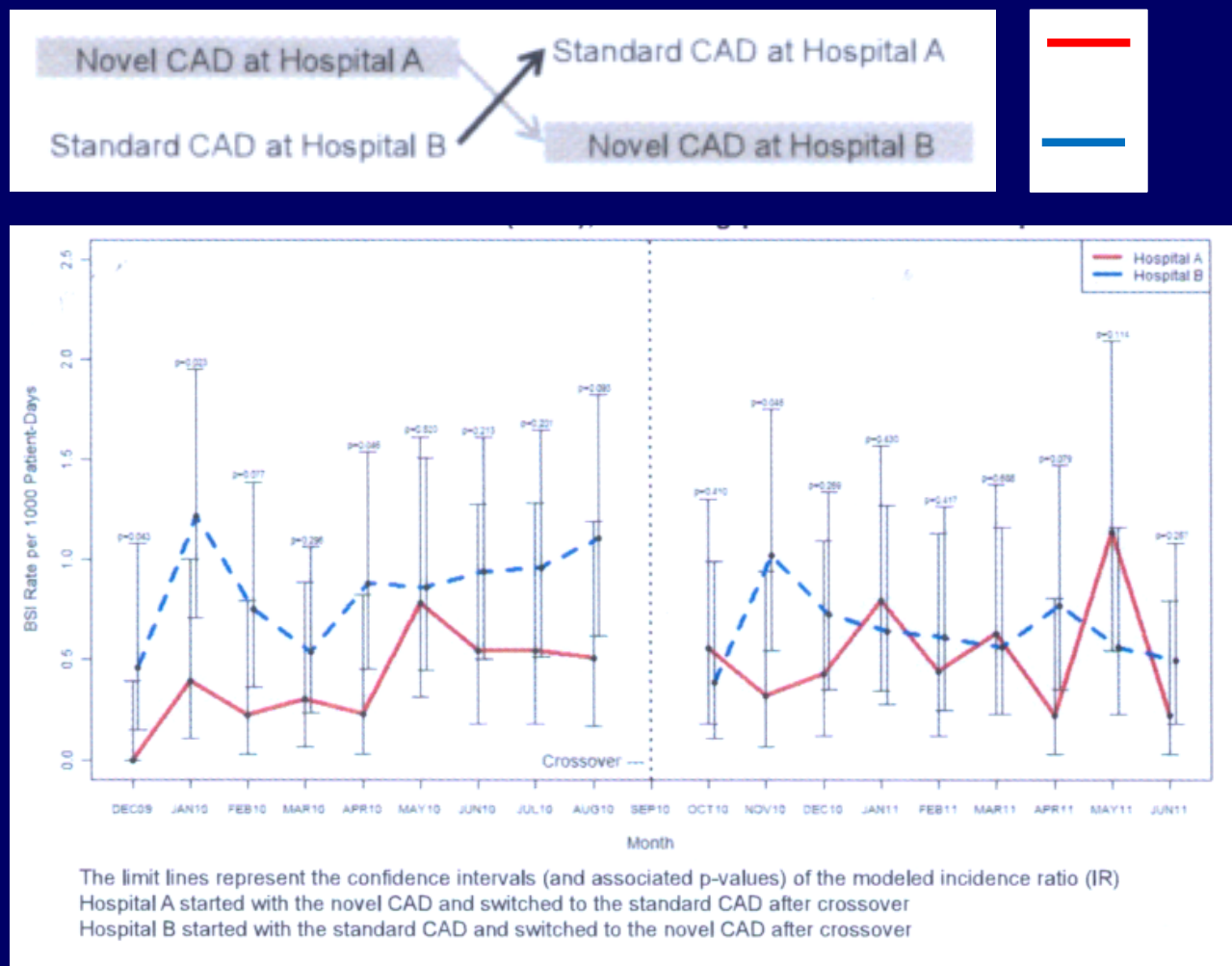
Silver coated connector valves



- 2 silver coated IV connector valves on the market. Very little clinical data re: effect on colonization of catheters or Bloodstream infection

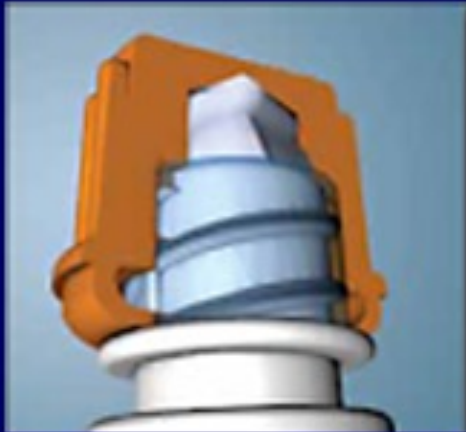
Comparison of a Novel Silver-Coated Needleless Connector and a Standard Needleless Connector for the Prevention Of CLA-BSI

JT Jacob, et al. ICHE, 2015



CLA-BSI rate:
1.79/1000 CVC d vs
1.21/1000 CVC d
IR = 0.68 [95% CI
0.52-0.89] P = 0.005

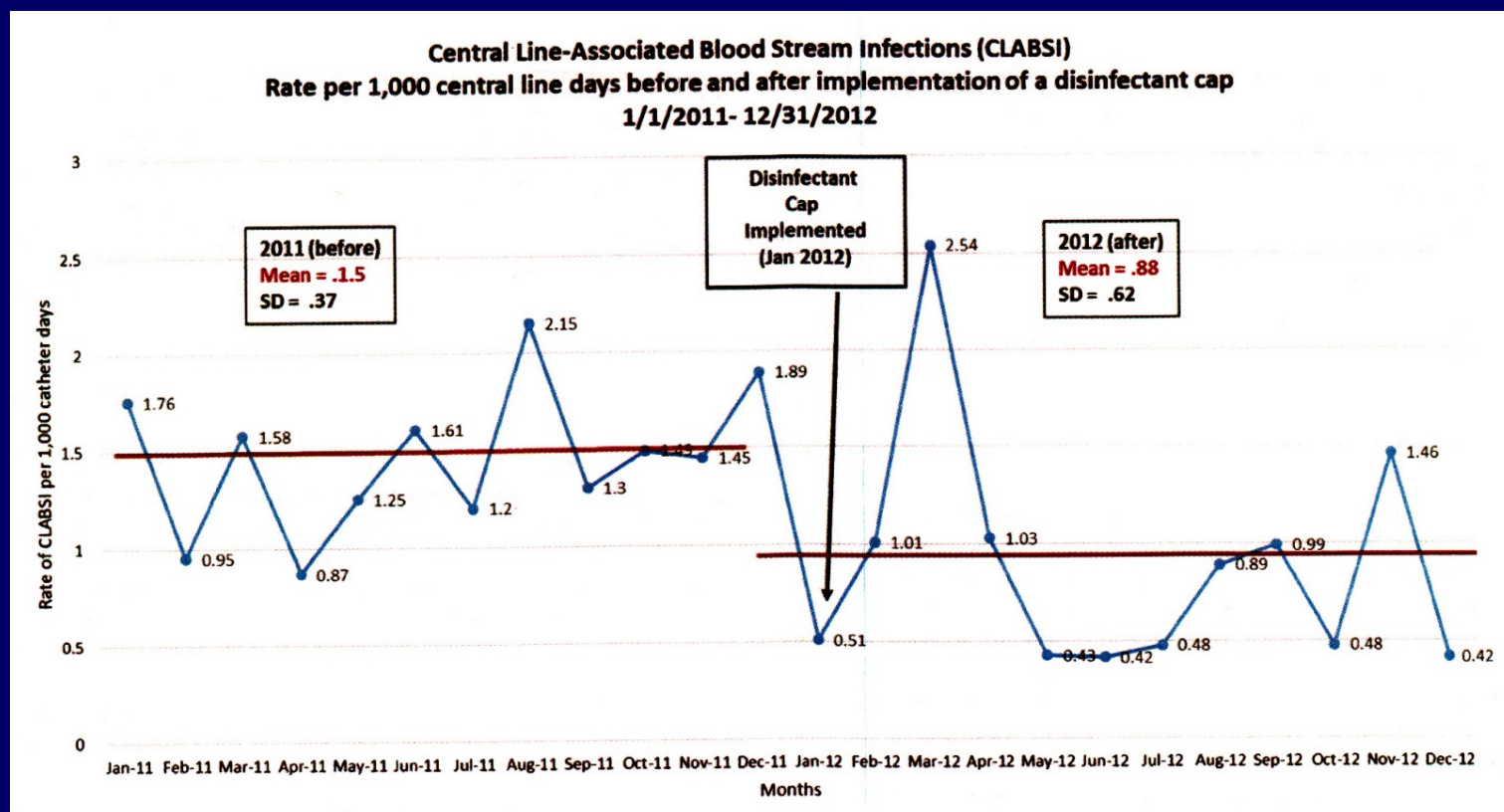
Antiseptic Caps



Impact of universal disinfectant cap implementation on central line–associated bloodstream infections

Katreena Collette Merrill RN, PhD^{a,*}, Sharon Sumner RN, BS^b,
Lorraine Linford RN, BS, CNSC^c, Carrie Taylor RN, MS, CIC^b,
Christopher Macintosh RN, BS^d

AJIC, 2014

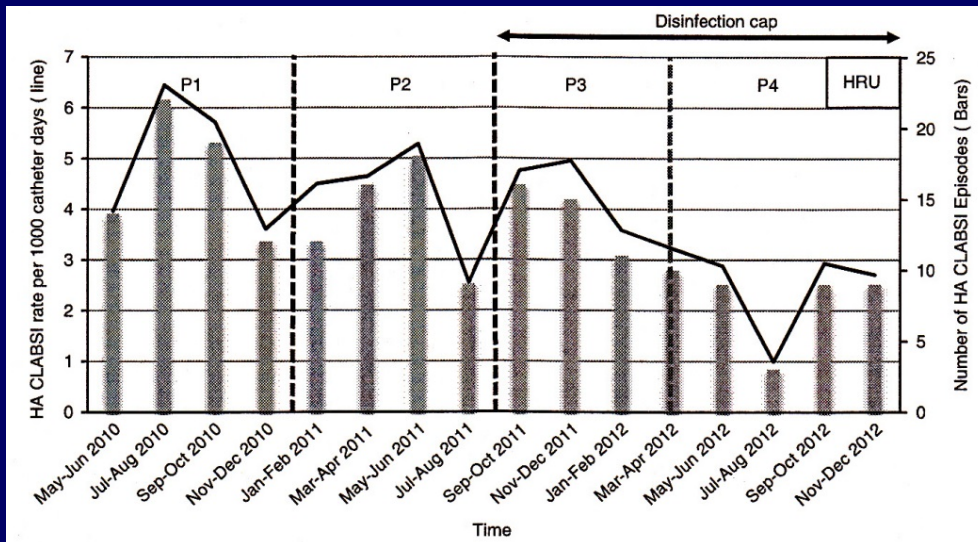


- CLA-BSI decreased from 1.5/1000 CVC d to 0.88/1000 CVC d, $p = 0.004$

Use of Disinfection Cap to Reduce Central-Line–Associated Bloodstream Infection and Blood Culture Contamination Among Hematology–Oncology Patients

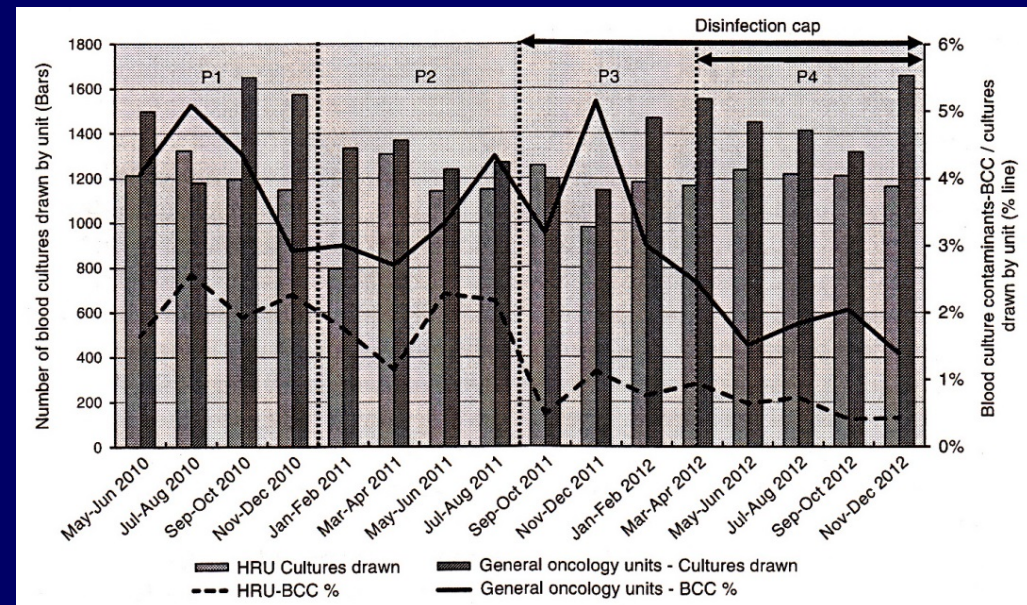
Mini Kamboj, MD;^{1,3,4} Rachel Blair, MPH;¹ Natalie Bell, RN;^{1,2} Crystal Son, MPH;¹ Yao-Ting Huang, MPH, PhD;³ Mary Dowling, MSN, RN;² Allison Lipitz-Snyderman, PhD;⁵ Janet Eagan, RN, MPH, CIC;¹ Kent Sepkowitz, MD^{1,3,4}

ICHE, 2015



CLA-BSI in high-risk pts: Rate per 1000 CVC d: 4.93, 4.22, 4.47, 2.34 (P1-P4 respectively)

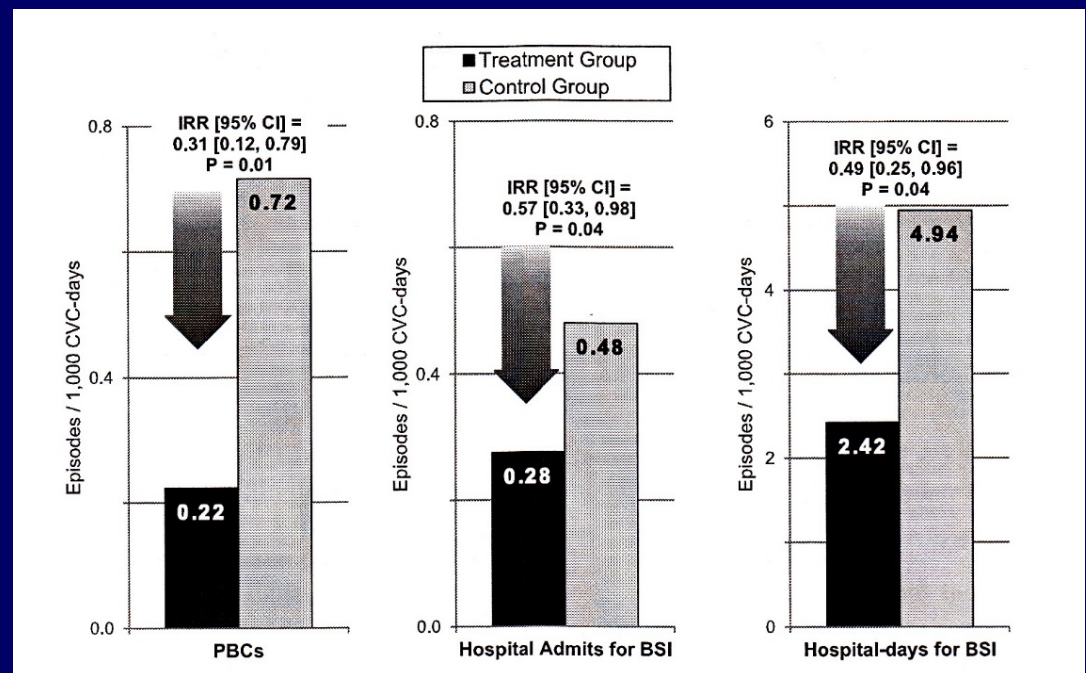
Blood culture contamination rate by CoNS



Dialysis Catheter–Related Bloodstream Infections: A Cluster-Randomized Trial of the ClearGuard HD Antimicrobial Barrier Cap

Jeffrey L. Hymes, MD,¹ Ann Mooney, MSN, RN, CNN,² Carly Van Zandt, MS,²
Laurie Lynch, PhD,³ Robert Ziebol, BS,³ and Douglas Killion, MBA³

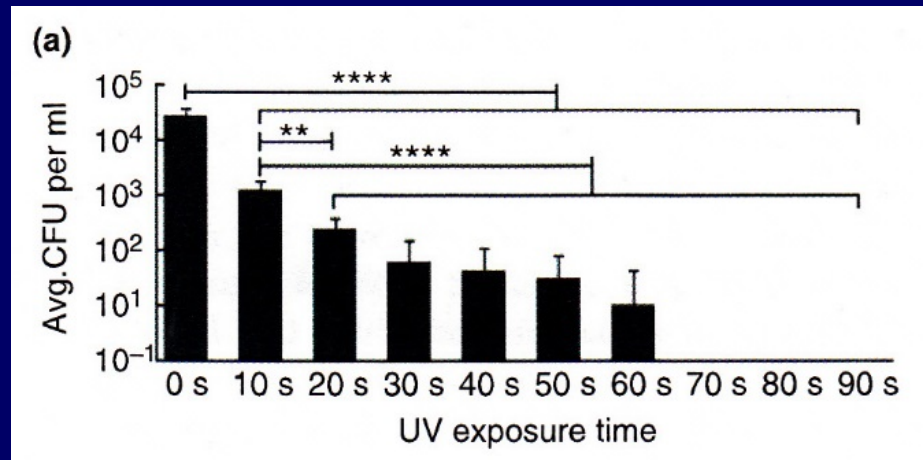
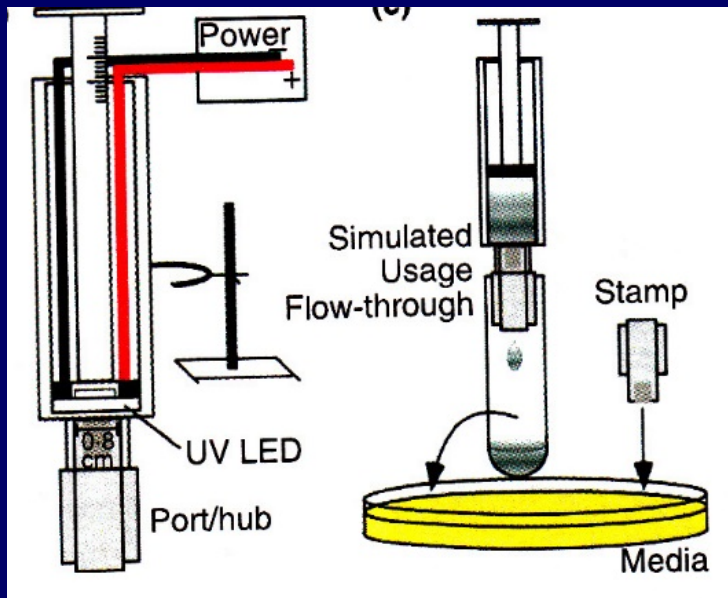
- 12 month, prospective, cluster-randomized study in 40 HD units.
- 2470 pts; 350,000 CVC days
- 56% lower (+) Bld Cx rate ($p=0.01$)
- 40% decrease in hospital admissions for BSI ($p=0.04$); 31% less hospital days ($p=NS$)



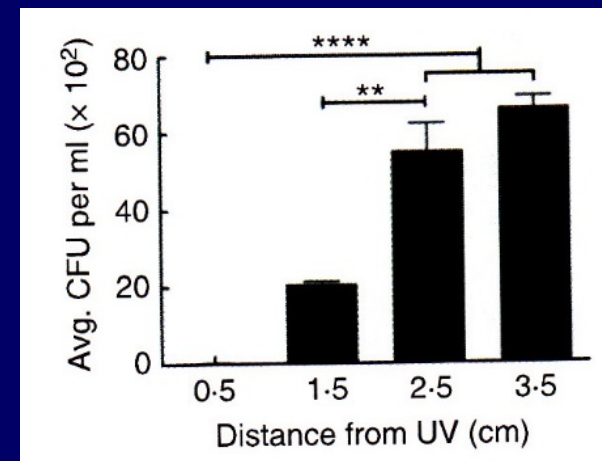
Results for last 6 months of the study

UV light-emitting diode disinfection

Hutchens, et al. J Applied Micro, 2015



- 285 nm UV LED effectively disinfected needleless connectors with 60s exposure at 0.5 cm.



Chlorhexidine Patient Bathing

Effectiveness of Chlorhexidine Bathing to Reduce Catheter-Associated Bloodstream Infections in Medical Intensive Care Unit Patients

Arch Intern Med 2007

Susan C. Bleasdale, MD; William E. Trick, MD; Ines M. Gonzalez, MD;
Rosie D. Lyles, MD; Mary K. Hayden, MD; Robert A. Weinstein, MD

- Daily CHG baths in ICU patients decreased BSI from 16.8 to 6.4/1000 CVC d.

Effect of Daily Chlorhexidine Bathing on Hospital-Acquired Infection

NEJM 2013

Michael W. Climo, M.D., Deborah S. Yokoe, M.D., M.P.H., David K. Warren, M.D.,
Trish M. Perl, M.D., Maureen Bolon, M.D., Loreen A. Herwaldt, M.D.,
Robert A. Weinstein, M.D., Kent A. Sepkowitz, M.D., John A. Jernigan, M.D.,
Kakotan Sanogo, M.S., and Edward S. Wong, M.D.

- 28% decrease in bloodstream infections ($P = 0.007$)

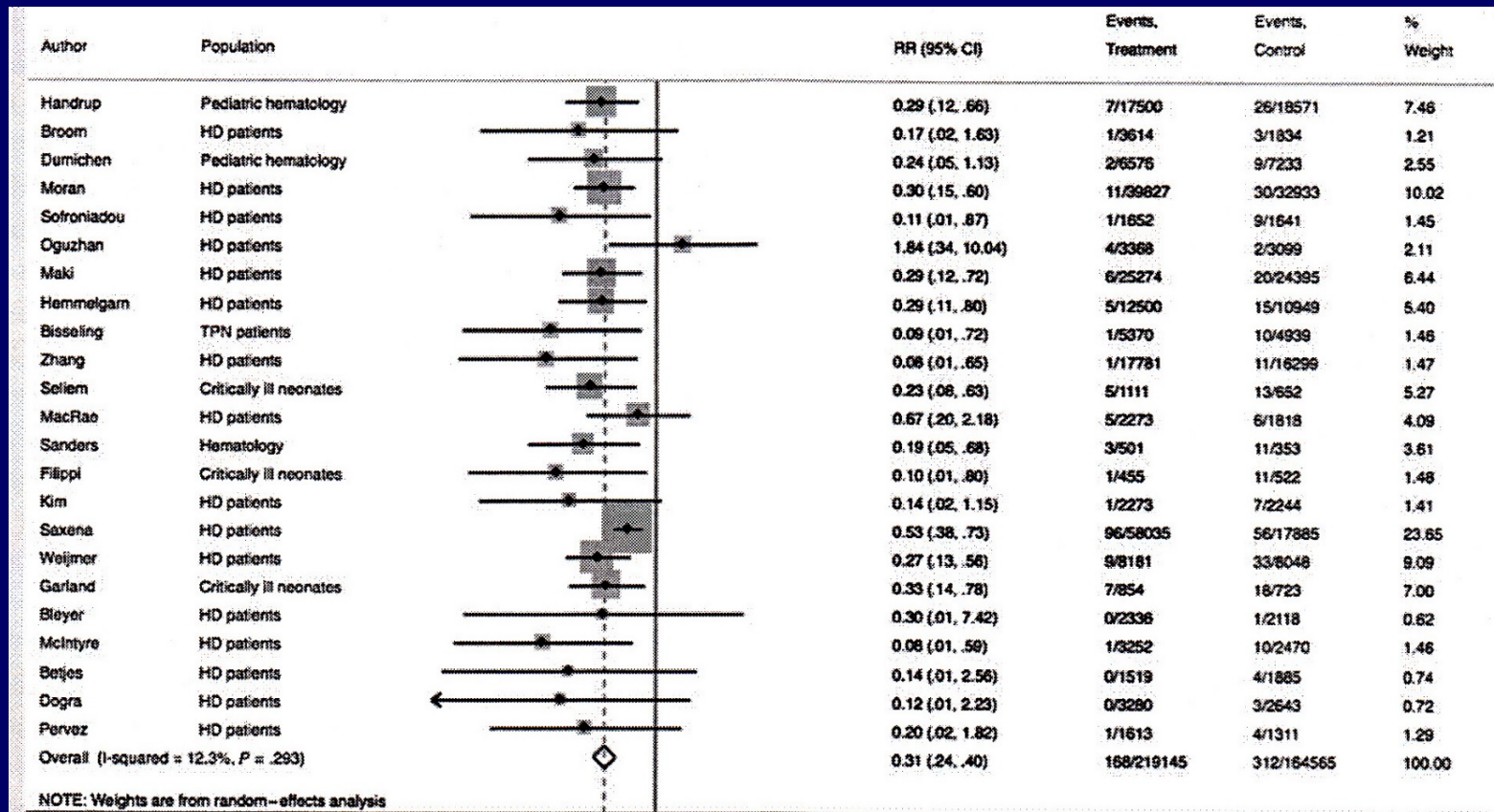
Targeted versus Universal Decolonization to Prevent ICU Infection

NEJM 2013

Susan S. Huang, M.D., M.P.H., Edward Septimus, M.D., Ken Kleinman, Sc.D., Julia Moody, M.S.,
Jason Hickok, M.B.A., R.N., Taliser R. Avery, M.S., Julie Lankiewicz, M.P.H., Adrijana Gombosev, B.S.,
Leah Terpstra, B.A., Fallon Hartford, M.S., Mary K. Hayden, M.D., John A. Jernigan, M.D., Robert A. Weinstein, M.D.,
Victoria J. Fraser, M.D., Katherine Haffenreffer, B.S., Eric Cui, B.S., Rebecca E. Kaganov, B.A., Karen Lolans, B.S.,
Jonathan B. Perlin, M.D., Ph.D., and Richard Platt, M.D., for the CDC Prevention Epicenters Program
and the AHRQ DECIDE Network and Healthcare-Associated Infections Program*

- HR for BSI (intervention vs baseline): 0.99 vs 0.78 vs 0.56 ($P = <0.001$)

Anti-Infective Catheter Lock Solutions



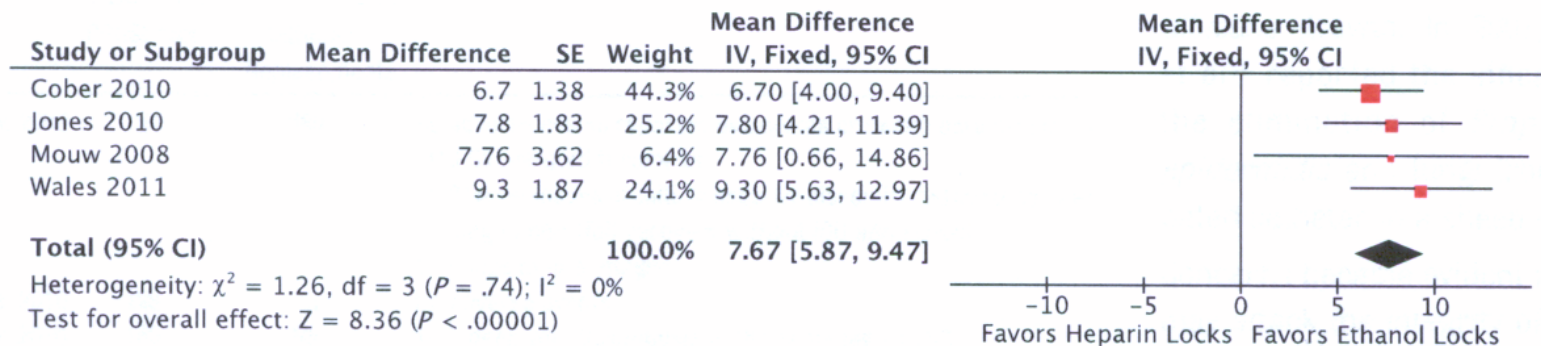
* 23 studies, 2896 patients, 69% decrease in CLABSI
RR: 0.31 95CI (.24-.40)
* 32% decrease in exit site infections
* Mortality 16% decrease (NS)

Conclusion: Anti-Infective Lock solutions are useful in certain circumstances. Additional study to assess optimal solution (antibiotics, alcohol, taurolidine, trisodium citrate, EDTA, nitroglycerin, etc) and populations

Ethanol Locks to Prevent Catheter-Related Bloodstream Infections in Parenteral Nutrition: A Meta-Analysis

Oliveira C, et al. Pediatrics, 2012

Pooled mean difference of CRBSI rate



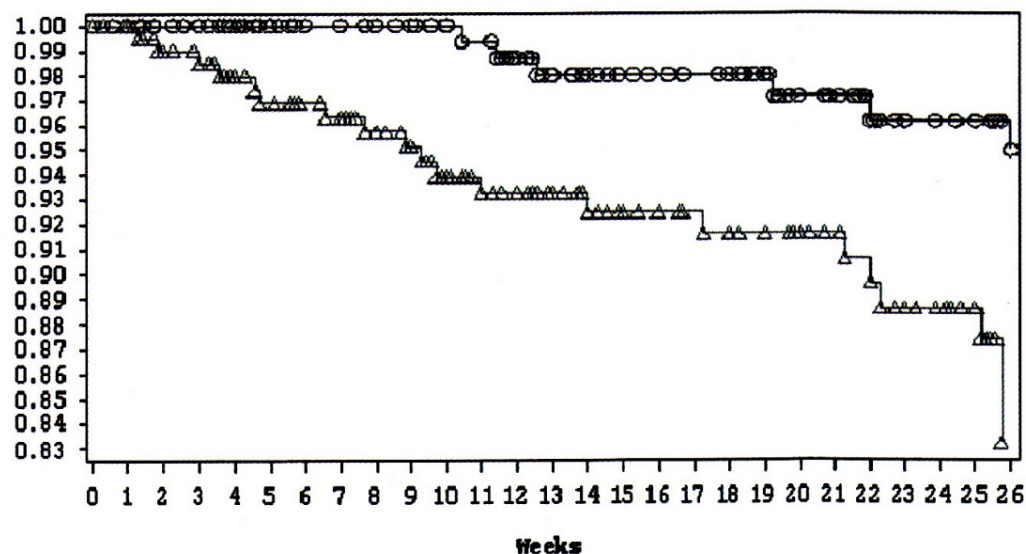
- Risk ratio for CR-BSI: 0.19 (95% CI 0.12-0.32)
- Risk ratio for catheter replacement: 0.28 (95% CI 0.06-1.23)
- Rare Toxicity: Etoh assoc with clotting, dizziness, CVC mechanical compromise, protein ppt, etc.

A novel antimicrobial and antithrombotic lock solution for hemodialysis catheters: A multi-center, controlled, randomized trial*

Dennis G. Maki, MD; Stephen R. Ash, MD; Roland K. Winger, BS, PE; Philip Lavin, PhD; for the AZEPTIC Trial Investigators

Crit Care Med 2011

The difference between the two groups is highly significant ($P=0.0016$ by log-rank test).



Weeks
 ○-○-○ C-MB-P △-△-△ Heparin

Patients remaining at risk by week

	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26
C-MB-P	201	198	190	185	177	171	164	162	160	157	152	150	143	138	132	124	121	119	118	114	107	103	97	92	90	89	82
Heparin	208	203	198	192	183	177	173	171	162	158	148	140	136	125	121	116	114	109	108	102	95	91	88	84	80	75	11

- Prospective, Randomized, Multi-Ctr trial
- 25 HD units, 407 pts, 50K CVC days
- 7% citrate, 0.15% methylene blue, 0.15% methylparaben, 0.015% propylparaben (C-MB-P)
- 0.82 vs 0.24 CRBSI/1000 CVC d; RR 0.29 (0.12-0.7, $p = 0.005$)

Statewide Survey of Technologic CLA-BSI Prevention

- Nebraska statewide survey of hospitals (25 PPS/65 CAH)
- Response: 17 PPS (68%), 25 CAH (40%)

Technology	PPS (%)	CAH (%)
CHG Dressing	94	73
Antibiotic or Antiseptic CVC	47	31
Passive port disinfection	35	54
CHG bathing in ICU	65	8
CVC lock soln	17	12

Scope of the Problem

What about Peripheral IVs???

Yearly Use of Peripheral IVs



- Little systematic data regarding complications: infection, phlebitis, infiltration, extravasation

**The Risk of Bloodstream Infection in Adults
With Different Intravascular Devices:
A Systematic Review of 200 Published Prospective Studies**

DENNIS G. MAKI, MD; DANIEL M. KLUGER, MD; AND CHRISTOPHER J. CRNICH, MD

Mayo Clin Proc, 2006

- Review of 110 studies, 10,910 catheters
- 0.1 BSI/100 devices; 0.5 (95% CI 0.2–0.7)/1000 device days
- 9 studies that required microbial concordance between catheter and blood culture: 0.6 BSI/1000 device d
- 1 per 1000 devices x 330 Million/2.25 attempts per successful IV start = 146,000 episodes of BSI

Status of Vascular Access at the University of Nebraska Medical Center

- Series of point prevalence surveys in all units during summer 2015
- All units visited on at least 3 occasions
- 755 patients
 - 59 (8%) No vascular access
 - 414 (55%) peripheral IV only
 - 239 (32%) CVC only
 - 43 (6%) both CVC and PIV

Peripheral IVs

Problems with
securement



Arterial Catheters & Dressings



Prevention of IV Catheter-Related Bloodstream Infection

■ Practice Measures

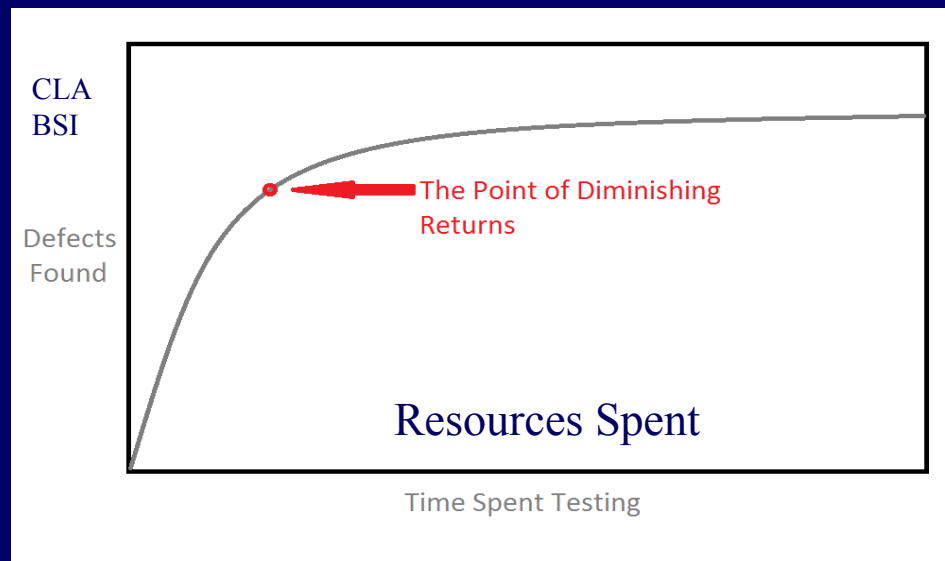
- Education and Training
- Appropriate staffing
- Insertion and Care Bundle
 - CHG skin prep
 - Sterile barrier precautions
 - Avoid femoral site
 - Scrub the hub
- Removal of CVCs

■ Technologic Innovations

- Antimicrobial-Coated CVC
- CHG impregnated dressings
- CHG patient bathing
- Catheter Flush/Lock soln
- Antimicrobial-Coated Connectors
- Antiseptic Caps

Prevention of IV Catheter-Related Bloodstream Infection and “Getting to Zero”

- Cost Effectiveness
- Point of Diminishing Returns



Questions & Comments

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HAND HYGIENE PROGRAMME AND ITS IMPACT ON ANTIMICROBIAL
RESISTANCE](#)

April 12, 2017

Speaker: **Professor Lindsay Grayson**, World Health Organization, Melbourne, Australia

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