# **PDR-Acinetobacter baumannii**: Can it be controlled?

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#### **Outlines**

Epidemiology of MDR-Acinetobacter baumannii in Thailand

> Is antimicrobial stewardship alone enough?

Implementing an Infection Control Program and rationale for each component

How to implement a successful infection control program in resource-limited settings?

#### National Resistance of A. baumannii



Source: Thai NIH

#### Percentage of Resistance of A. baumannii complex



#### Risk factors and Outcomes of Carbapenem Resistance *A. baumannii*

- > Increasing 3<sup>rd</sup> GC/ carbapenem consumptions.
- > Patients being transferred from another hospital.
- > Duration of 3<sup>rd</sup> GC/ carbapenem use.
- > Environmental contamination.
- Mortality rate 38% among carbapenem resistant A. baumannii (CRAB) vs. 24% carbapenem sensitive A. baumannii (CSAB) (95% CI, 1.2-2.2).
- > BSI is a predictor for mortality.
- Carbapenem resistance associated with higher antibiotic costs and admission costs.

Apisarnthanarak A, et al. Predictors for mortality among *PDR-A. baumannii* infections in Thailand. Am J Infect Control, 2009; Jamulitrat S, et al. An outbreak of imipenem-resistant Acinetobacter baumannii at Songklanagarind Hospital: the risk factors and patient prognosis. J Med Assoc Thai, 2007.; Surasarang K, et al. Risk factors for multi-drug resistant *Acinetobacter baumannii* nosocomial infection. J Med Assoc Thai, 2007.

#### Antimicrobial Stewardship at Thammasat University Hospital

#### > Educations to intern, residents, and staff

- Five-sessions education for all intern, residents and staff in the hospital
- One on one feedback to externs, interns, and residents

#### Intervention target at specific units

- Structured teaching on medicine, surgery, and OB-GYN
- Monthly education for medical students, interns, and residents
- Presence of clinical pharmacist on medicine, surgery and OB-GYN units

#### Encourage Infectious Diseases consultation Initiated in July 2003

Apisarnthanarak A, et al. Effectiveness of Educational and Antibiotic Control Program in a tertiary care hospital in Thailand, Clin Infect Dis, 2006

#### **Bacterial Resistance Rates During the Pre and Post Intervention Periods**

	Resistance rate % <sup>a</sup>					
Microorganism	Pre intervention period	Post intervention period	Associated antibiotic class	Type of variation	R <sup>b</sup>	Р
MRSA	48	33.5	Glycopeptides	Decrease	0.55	<.001
			3 <sup>rd</sup> gen. Cephalo.	Decrease	0.93	<.001
ESBL <i>E.coli</i>	33	21	3 <sup>rd</sup> gen. Cephalo.	Decrease	0.74	<.001
ESBL K.pneumoniae	30	20	3 <sup>rd</sup> gen. Cephalo.	Decrease	0.69	<.001
3 <sup>rd</sup> GC-resistant <i>A. baumanii</i>	27	19	3 <sup>rd</sup> gen. Cephalo.	Decrease	0.78	<.001
Imipenem-resis <i>P. aeruginosa</i>	5	4	None	-	-	
MDR <i>A.baumanii</i>	4	5	None	-	-	

<sup>a</sup> Calculated using the total number of strains.

<sup>b</sup> Linear regression analysis between evolution in the resistance rate and antibiotic use throughout the study

#### Controlling Antibiotic Use and Resistance (Editorial Note)

".....To sustain or even further improve these results, lasting and repeated efforts will be needed. Integrating infection-control efforts into this education and antibiotic-control program is warranted."

Nouven JL, Clin Infect Dis, 2006

# The emergence of *PDR-A. baumannii* outbreak: The need to emphasize on infection control program in Thailand

## Despite effective antimicrobial stewardship program, *PDR-A. baumannii* emerged....



### What works!

- Most outbreaks were terminated with multi-faceted, comprehensive infection control programs
- Measures always include education, hand hygiene (5moments), contact isolation, environmental cleaning, targeted active surveillance culture in high risk area, and antimicrobial control program
- Recent reports also suggested the role of 4% chlorhexidine total body wash

Dancer SJ. JHI 2009 Rodriguez-Bano J, AJIC 2009 Valencia R, ICHE 2009 Gill CJ, CID 2009 Borer A, JHI 2007 Chan PC, ICHE 2007

#### Controlling healthcare associated Infection: Vertical vs. horizontal approach





A. Borer<sup>a,\*,</sup> J. Gilad<sup>a</sup>, N. Porat<sup>b</sup>, R. Megrelesvilli<sup>c</sup>, L. Saidel-Odes <sup>a</sup>, N. Peled<sup>d</sup>, S. Eskira<sup>a</sup>, F. Schlaeffer<sup>e</sup>, Y. Almong<sup>c</sup>



Figure 2 Patient population changes and *Acinetobacter baumannii* (ACBA) skin colonisation by time of culture. In-ward population: colonised and non-colonised ACBA patient population in the medical intensive care unit. Outof-ward population: patients discharged from the unit either colonised or non-colonised at time of discharge.

### **Use of Hydrogen Peroxide Vapour**

Infect Control Hosp Epidemiol. 2010 Dec;31(12):1236-41. Epub 2010 Oct 25.

#### Use of vaporized hydrogen peroxide decontami Acinetobacter baumannii infection at a long-ter

Ray A, Perez F, Beltramini AM, Jakubowycz M, Dimick P, Jacobs MR, Roman K, Bonon **OBJECTIVES:** To describe vaporized hydrogen peroxide (VHP) as an adjuvant in the conspital (LTACH) and to describe the risk factors for acquisition of MDR A. baumannii in

DESIGN: Outbreak investigation, case-control study, and before-after intervention trial.

SETTING: A 54-bed LTACH affiliated with a tertiary care center in northeastern Ohio.

METHODS: Investigation of outbreak with clinical and environmental cultures, antimicrol type strains, and case-control study; and intervention consisting of comprehensive infect

RESULTS: Thirteen patients infected or colonized with MDR A. baumannii were identifie carbapenem-resistant. MDR A. baumannii was found in wound samples, blood, sputum, associated pneumonia was the most common clinical syndrome caused by the pathoger found in 8 of 93 environmental samples, including patient rooms and a wound care cart; immediately after VHP decontamination and both 24 hours and 1 week after VHP decon When patients colonized with MDR A. baumannii reoccupied rooms, environmental contamination recurred.

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**CONCLUSION:** Environmental decontamination using VHP combined with comprehensive infection control measures interrupted nosocomial transmission of MDR A. baumannii warrants further investigation.

### Implementation of IC Program

- Active surveillance (tracheal aspirate and/or rectal swab) in ICU patients.
- Repeat surveillance cultures weekly.
- Cohort and contact isolation.
- Hand hygiene intervention
- > Environmental cleaning
- Monitoring of Infection Control adherence

### Gown change every shift

มีระจำเสี่			INSAT		B JIH MARINA		
ประเภทบุคลากรการแพทย์	ก่อน	หลัง	ข้อสังเกตุจากการเฝ้าระ รักกละของการเข้าสะเมืองเป็น	ประเภทบุคลากรการแพทย์	ก่อน	หลัง	ข้อสังเกตุจากการเป้า
แพทย์	0.00 %	16.67 %		แพทย์	43 % (7)	85% (7)	
พยาบาล	0.00%	40.00%		พยาบาล	10 % (10)	60% (10)	
เจ้าหน้าที่อื่นๆ	0.00 %	0.00%		เจ้าหน้าที่อื่นๆ	17% (6)	67 % (6)	
รวม	0.00 %	12.50 %		รวม	22 % (23)	70 % (23)	-

2

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### Hand Hygiene Compliance Rate



#### Characteristics of patients in intervention units during the entire study periods (Cont.)

Characteristics	Period 1	Period 2	Period 3
	n= 1,357	n= 1,273	n= 1,441
Source of A. baumannii infection and colonization			
Bloodstream	14 (27)	4 (24)	3 (23)
Urinary tract	4 (7)	1 (6)	1 (8)
Pulmonary	31 (59)	11 (64)	8 (61)
Other	4 (7)	1 (6)	1 (8)
Admissions rate (mean admission per day)	3.7	3.5	3.9
APACHE-II score, mean±SD	17 ± 5	17 ±4	18 ±5
Total number of patient-day	14,650	14,456	15,410
Daily occupancy, mean days $\pm$ SD	$20.2 \pm 1.4$	$21 \pm 2.4$	$20 \pm 2.4$
Duration of hospital stay, mean days $\pm$ SD	$10.4 \pm 4.7$	$11.5 \pm 3.5$	$10.6 \pm 3.5$
Patients with contact isolation,	8.8 ±1.9	5.5 ±2.1	3.8 ±1.2
mean patients/d ±SD			

#### Patient colonized or infected with *PDR-Acinetobacter baumannii*, infection control compliance monitoring, and outcomes in intervention units

Variable	Period 1	Period 2	Period 3
Environmental cleaning			
No. of observations		166	165
Mean no. of observations per week $\pm$ SD		$3 \pm 0.6$	$3 \pm 0.4$
Environmental cleaning rate, mean $\pm$ SD		$0.85 \pm 0.08$	$0.83 \pm 0.09$
Hand hygiene adherence			
No. of observations	154	166	165
Hand hygiene adherence rate before	$0.31 \pm 0.07$	$0.75 \pm 0.08$	$0.54 \pm 0.01$
and after contact, mean $\pm$ SD			
Hand hygiene adherence rate before	$0.24 \pm 0.02$	$0.63 \pm 0.02$	$0.51 \pm 0.09$
and after contact and glove&gown use,			
mean±SD			

#### Patient colonized or infected with *PDR-Acinetobacter baumannii*, infection control compliance monitoring, and outcomes in intervention units

Variable	Period 1	Period 2	Period 3
Outcomes			
Rate of PDR- Acinetobacter baumannii acquisitions, isolate per 1000 patient-		15.9	11.9
days at-risk			
Rate of PDR- <i>Acinetobacter baumannii</i> infection and colonization (/1000 patient-days)	3.6	1.2	0.85
Monthly antibiotic cost for PDR- A. baumannii treatment (USD)	3,762±605	1,722±96	1,278±87
Hospitalization cost for each patient (USD)	366±100	252±96	204±88

### **Cost-Benefit of the Interventions**

> Total cost for ASCs \$19,862.

Compared to period 1, the monthly ATB cost reduced by 36-42% in period 2 and 3. (\$3,762 vs. \$1,776 vs. \$1,278)

Compared to period 1, hospitalization cost reduced by 25-36% in period 2 and 3. (\$366 vs. \$254 vs. \$204)

Apisarnthanarak A, et al. A multi-faceted infection control intervention to reduce *PDR-A. baumannii* in three ICUs in a Thai tertiary care center. Clin Infect Dis, 2008

#### **Concurrent Initiative to Reduce Nosocomial Infections**

### **Urinary Tract Infection Intervention**



Apisarnthanarak A, et al. Effectiveness of multifaceted hospital wide quality improvement program featuring intervention to remove IUC in a tertiary care center in Thailand. ICHE, 2007

#### **Implementing VAP Bundle**



#### **Implementing CA-BSI Bundle**



Apisarnthanarak A, et al. Effectiveness of CA-BSI bundle. AJIC, 2010

#### Impact of Concurrent Interventions to Reduce Healthcare-Associated Infection

Characteristic	Period I (n = 1115)	Period I (n = 1115) Period 2 (n = 1050)	
Organism associated with CA-BSI* <sup>,‡</sup>			
Methicillin-susceptible Staphylococcus aureus	21 (24)	8 (25)	2 (28)
Methicillin-resistant S aureus	18 (20)	7 (21)	2 (28)
Coagulase-negative Staphylococcus spp	26 (30)	10 (32)	2 (28)
MDR-Acinetobacter baumannii*	23 (23)	4 (I2) <sup>†</sup>	l (14)†
Pseudomonas aeruginosa	7 (15)	2 (7)	0 (0) <sup>†</sup>
ESBL-producing Escherichia coli	7 (8)	3 (9)	0 (0) <sup>†</sup>
ESBL-producing Klebsiella pneumoniae	4 (5)	I (5)	0 (0) <sup>†</sup>

Intervention to reduce NI clearly impacted the incidence of *MDR-A. buamannii* within 1 year

### After the intervention.....



Apisarnthanarak A, et al. A multi-faceted infection control intervention to reduce *PDR-A. baumannii* in three ICUs in a Thai tertiary care center. Clin Infect Dis, 2008

### Why Active Surveillance?



#### Which is the most appropriate site for surveillance A. baumannii?

JOURNAL OF CLINICAL MICROBIOLOGY, Nov. 1997, p. 2819-2825 0095 - 1137 / 97 / \$04.00 + 0Copyright © 1997, American Society for Microbiology

#### Distribution of *Acinetobacter* Species on Human Skin: Comparison of Phenotypic and Genotypic Identification Methods

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				- membranes of patients and controls				
	No. (%) of patients or controls colonized with <i>Acinetobacter</i> spp.			memoranes of patients and controls				
Body site				Genomic species	No. (%) of Acinetobacter strains isolated from:			
	Patients	Controls	groups	(DNA group) <sup>a</sup>	Patients	Controls	Both	
orehead	13 (33)	5(13)	18 (23)				groups	
ar	14 (35)	3 (8)	17 (21	A. baumannii (2) <sup>b</sup>	2(1)	1 (3)	3 (1.5)	
ose	13 (33)	3 (8)	16 (20	Acinetobacter sp. 3	18(12)	26	20 (11)	
hroat	6 (15)	0 (0)	6 (8)	A. junii (5)	6 (4)	3 (10)	9 (5)	
xilla	13 (33)	1 (3)	14 (18	A. johnsonii (7)	34 (22)	6 (20)	40 (21)	
and	13 (33)	8 (20)	21 (26	A. lwoffii (8/9)	69 (44)	18 (58)	87 (47)	
roin	15 (38)	5 (13)	20 (25	Acinetobacter sp. 10	$1(1)^{'}$	ര്ത്	1(0.5)	
erineum	8 (20)	1 (3)	9 (11	A. radioresistens (12)	22 (14)	0 (0)	22 (12)	
oe web	16 (40)	3 (8)	19 (24	Unidentified	3 (2)	1 (3)	4 (2)	
Total no. (%) of sites colonized	111 (31)	29 (8)	140 (19	Total no. of isolates	155	31	186	

TABLE 2. Rates of colonization of various body sites of patients and controls with Acinetobacter spp.

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TABLE 1. Distribution of different Acinetobacter spp. identified by phenotypic properties after isolation from the skin and mucous

Vol. 35, No.

#### The Inanimate Environment Can Facilitate Transmission

#### X represents *A. baumannii* culture positive sites



~ Contaminated surfaces increase cross-transmission ~

#### "Role of environmental cleaning"



Patient with MDR-A. baumannii

### Interactions

То	<b>Interventions</b>
Environment	НН
HCW's Hand	НН
	Environment cleaning
Patient	НН
	Contact Isolation
HCWs Hand	НН
	Contact isolation
	Environment cleaning
	To Environment HCW's Hand HCWs Hand

### Monitoring for 5 Moment Hand Hygiene



# Main Menu Email Observations



**Reset:** 



Send

#### Tap to Resend Email

**Email Observations:** 

2 Observations

### Why Sodium Hypochlorite?

- Several studies have utilized sodium hypochlorite to control MDR-A. baumannii outbreak successfully.
- Floor cleaning with detergent did not effect the NI rates.
- Quaternary ammonium compound was reported to be inadequate to clean bathroom and toilets.

Denton M, et al. JHP, 2004 Pimental JD, et al. JHP, 2005 Das I, et al. JHP, 2002 Danforth D, et al. JHP, 2007

#### **Adverse Effect from Sodium Hypochlorite**

#### Damage HCW's skin



#### Damage hospital floor



#### Descending Order of Resistance to Germicidal Chemicals

#### **Bacterial spores**

*B. subtilis* Clostridium sporogenes

#### **Mycobacteria**

M. tuberculosis var. bovis

#### Non-lipid or small viruses

Polio virus, rhinovirus, norovirus

#### Fungi

Trichophyton, Candida Cryptococcus

#### **Vegetative bacteria**

Pseudomonas, staphylococci (MRSA), enterococci (VRE)

#### Lipid or medium-sized viruses

HBV, HIV, HCV, HSV, CMV, Ebola



# Creating cohort area to limit transmission of *PDR-A. baumannii* in a medical unit

- October 2007, first case of PDR-A. baumannii was detected in a medical unit. The nurse to patient ratio was 1: 8 in this medical unit.
- IC measured were implemented within 24 hours including 1) enhanced contact isolation, 2) ASCs, 3) environmental cleaning, 3) enhanced hand hygiene program
- During period 1 (4-28 October), 6 cases of PDR-A. baumanii were detected by ASCs; infection and colonization rate 2.4/1000 patient-days & acquisition rate 6/1000 patient-days.

Apisarnthanarak A, et al. Creating cohort area to limit transmission of *PDR-A. baumannii* in a medical unit. CID, 2009 40

#### **Developing IC Intervention Need to Understanding Human Behavior**

### **VDO** Clips

### **Toilet without Intervention**



#### **Toilet with Intervention**



### **Toilet with Intervention**



Will you wash you hand, if you see your hand like these pictures?



# We need to have faith for believing that it is possible to IC Compliance

"Positive Deviance" VDO Clip



initially presented at the International Quality Symposium, Mexico city, 1989. It indicated how management's failure to understand its processes and practices from the perspective of its customers, suppressed the company's profits by as much as 40%.

<u>Front line workers</u>: experts at the work they do, decide HOW to do work, & foster self-discovery among peers; <u>owners</u>

Leadership and middle managers support and filter ideas, and remove barriers for implementation of practices from frontline workers

PD

48

### Steps for PD implementation

1: Don't presume you have the answer. 2: Don't think of it as a dinner party. 3: Let them do it themselves. 4: Identify conventional wisdom. 5: Identify and analyze the deviants. 6: Let the deviants adopt deviations on their own.

- 7: Track results and publicize them.
- 8: Repeat steps 1 through 7.

### **Positive deviance (PD)**

"PD program was implemented by identifying colleagues that were highly compliant with hand hygiene. All HCWs attended PD meetings. At these meetings, we have representative HCW from each shift. During the meetings the PD discuss problems that they notice (e.g., Dr. X did not handwash their hands during the patient examination). We discuss ways to stimulate talking to noncompliant individuals in a positive manner. No humiliation is permitted. The positive deviants are also given the opportunity to express their feeling, to discuss among them what needs to be changed, what needs to be improved, what is to be taken as a good example. We encourage them to invite another positive deviant to the next meeting." Alex Marra, MD









# Creating cohort area to limit transmission of *PDR-A. baumannii*

During period 2 (November 07-January 08): a cohort area was created in an assigned section, educational session on HH adherence and realtime feedback on IC processes. One nurse/shift was assigned to the cohort area.

The infection and colonization rate 0/1000 patientdays (P = 0.05). PDR-A. baumannii acquisition rate was 1/1000 patient-days (P < 0.001).</p>

Apisarnthanarak A, et al. Creating cohort area to limit transmission of *PDR-A. baumannii* in a medical unit. CID, 2009



In January 2005 Thammasat University Hospital, a 500-bed teaching hospital in central Thailand, began a three-year project to eliminate pandrug-resistant *Acinetobacter baumannii* (PDRAB), a Gram-negative bacteria intrinsically resistant to most antibiotics that is a significant source of HAIs in Thailand, particularly in ICUs. After an initial data analysis, a team led by Anucha Apisarnthanarak, MD, devised an action plan in the hospital's three ICUs consisting of the following:

- Gowns and gloves used during all patient care, plus scrupulous hand hygiene both before and after patient contact in all ICUs;
- Screening for PDRAB upon ICU admission and every seven days until discharge;
- Cohorting of colonized or infected patients; and
- Sanitizing surfaces with 1:100 solution of sodium hypochlorite (later changed to detergents and ammonia due to the corrosive effect on skin and surfaces).

The hospital encouraged compliance using educational sessions, wall posters, and monthly feedback to staff. Infection control specialists monitored both caregivers and housekeepers as they went about their jobs. The result: four months — two of them consecutive — with zero colonization or infection. Overall, the rate was reduced from 3.6 cases per 1,000 patient days before the intervention to 0.85 cases at the end of 2007.

Since then, Apisarnthanarak and his team have continued the protocol, minus the active screening, and infection rates have remained stable. The effort, he says, shows that "in a resource-limited setting where there is not much technology, simple and easily implemented infection control can reduce even the most drug resistant microorganism."

# In which settings do these strategies work?



These strategies might not work in hospitals settings that cannot follow Ningtingale's suggestion.

Corbis.com

#### **Examples of those settings**



#### What to do to Control *MDR-Acinetobacter* in Resource-Limited Settings?

#### **Resource-full**

- Molecular epidemiology
- > Environmental culture
- Active Surveillance
- Enhanced environmental cleaning
- Enhanced isolation precaution
- > Antibiotic management

#### **Resource-Limited**

- Stratified unit specific infection rate
- Line listing and/or case-control study (identify common source outbreak)
- Implement emergency measure for highly alert pathogen
- Initial environmental culture (per finding from line listing)
- Modified ASC
- > Enhanced isolation precaution
- Environmental cleaning
- Antibiotic management program

### Conclusions

- Infection Control intervention is crucial to help reduce MDRmicroorganisms in developing country.
- Infection Control component must be modified to fit each setting & infrastructure.
- Integration of multi-faceted infection control program can successfully reduce MDR-microorganisms.
- Monitoring of adherence to infection control component is important to help sustain reduction of MDR-microorganisms.
- Additional studies on infection control interventions and behavior science to reduce MDR-microorganisms are needed.

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### Thank you for your attention

