

Welcome to *California*



# Carbapenem-Resistant Enterobacteriaceae (CRE) in California

Greater Los Angeles APIC  
Los Angeles, CA  
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# Learning Objectives

- Review CDC Toolkit core CRE prevention strategies for ALL healthcare facilities
- Describe the background, methods, and results of the statewide CRE 2012 Prevalence Survey with a focus on Los Angeles County
- Describe CRE control in regions where there are few CRE and discuss guidance for regions where CRE are common

*Note: No regions in CA meet definition of "Common" in 2012*

# Carbapenem-resistant *Enterobacteriaceae* (CRE)

- One of three pathogens on CDC's most urgent threat list



- CRE infections result in 40-50% mortality
- CRE carry genes that are transmissible and confer resistance to many other antimicrobials, leaving limited treatment options

- Pan-resistant CRE *Klebsiella* have occurred

# Definitions of CRE

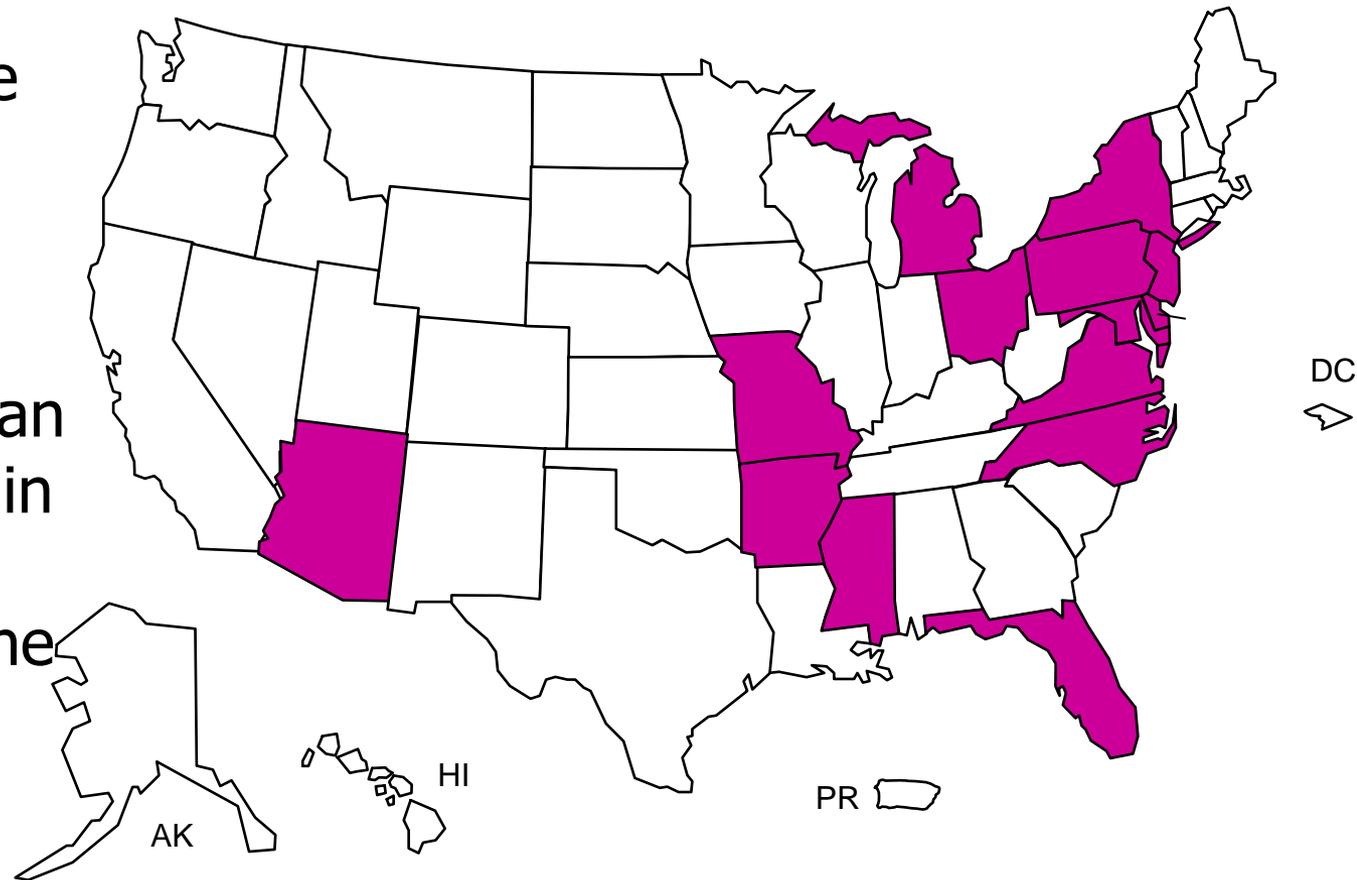
- Nonsusceptible to **one** of the following carbapenems:  
doripenem, meropenem, or imipenem

**- AND -**

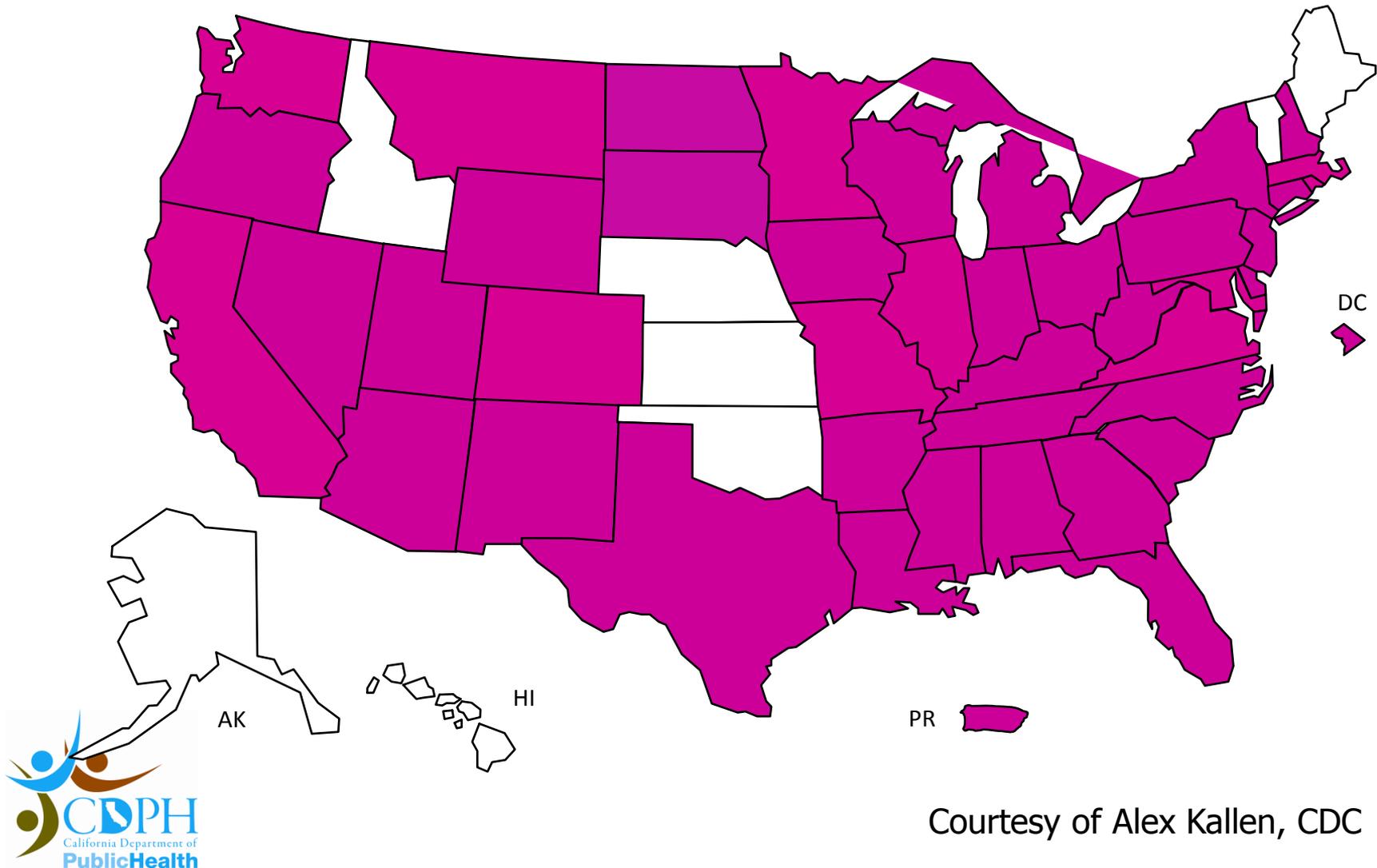
- Resistant to **all 3** of the following 3<sup>rd</sup> generation cephalosporins (or all that were tested):  
ceftriaxone, cefotaxime, and ceftazidime
- Note: All three of these antimicrobials are recommended as part of the primary or secondary susceptibility panels for *Enterobacteriaceae* testing

# Emergence of CRE in United States

- CRE was first identified in the US in 1996 in North Carolina
- By November 2006, CRE began to be reported in a number of states across the country



# KPC-Producing CRE in US, 2013

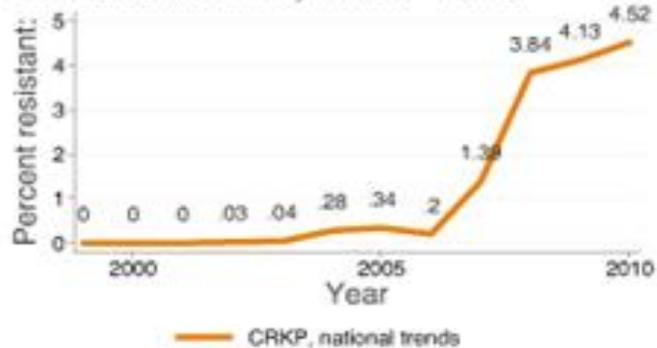


# Carbapenam-resistant *Klebsiella pneumoniae*

US Regions, 2009-2010



National trends, 1999-2010



ResistanceMap

[cddep.org/map](http://cddep.org/map)

# CRE Risk Factor: Care in LTAC Hospitals

- A Chicago survey found patients in LTAC hospitals had a significantly greater risk of being colonized or infected with CRE than patients in short term stay hospitals
  - RR= 5.94, 95% CI: 3.75-9.39
- In an outbreak of 40 CRE cases in Indiana and Illinois, 24 (60%) were linked to one LTAC hospital



Lin M et al. Clin Infect Dis. 2013;57:1246-1252  
Won S Y et al. Clin Infect Dis. 2011;53:532-540

# Other CRE Risk Factors

- CRE infection or colonization – New York data
  - Exposure to cephalosporins (OR: 2.65,  $p=.02$ )
  - Exposure to carbapenems (OR: 14.97,  $p<.01$ )
  - Transplant (OR: 3.71,  $p=.008$ )
  - Pre-Infection LOS (OR: 1.05,  $p=.01$ )
  - Ventilator (OR: 2.44,  $p=.04$ )
- CRE infection – Israel data
  - Poor functional status (OR: 15.4,  $p<.01$ )
  - ICU stay (OR: 17.41,  $p=.02$ )
  - Receipt of antibiotics (OR: 4.4,  $p=.05$ )
    - Fluoroquinolones (OR: 7.2,  $p=.04$ )

## Primary Prevention? **CRE Surveillance!**

- Every inpatient healthcare facility needs to have awareness of whether CRE (*E.coli* or *Klebsiella*) have ever been cultured from admitted patients
- Facilities without this information should review archived lab results from previous 6 months or a year
- If CRE have been present, determine
  - If evidence of intra-facility transmission
  - Which units/wards affected
  - Basic epidemiology of CRE patients, including dates of admission, clinical outcomes, medications, common exposures (i.e. wards, surgery, procedures)

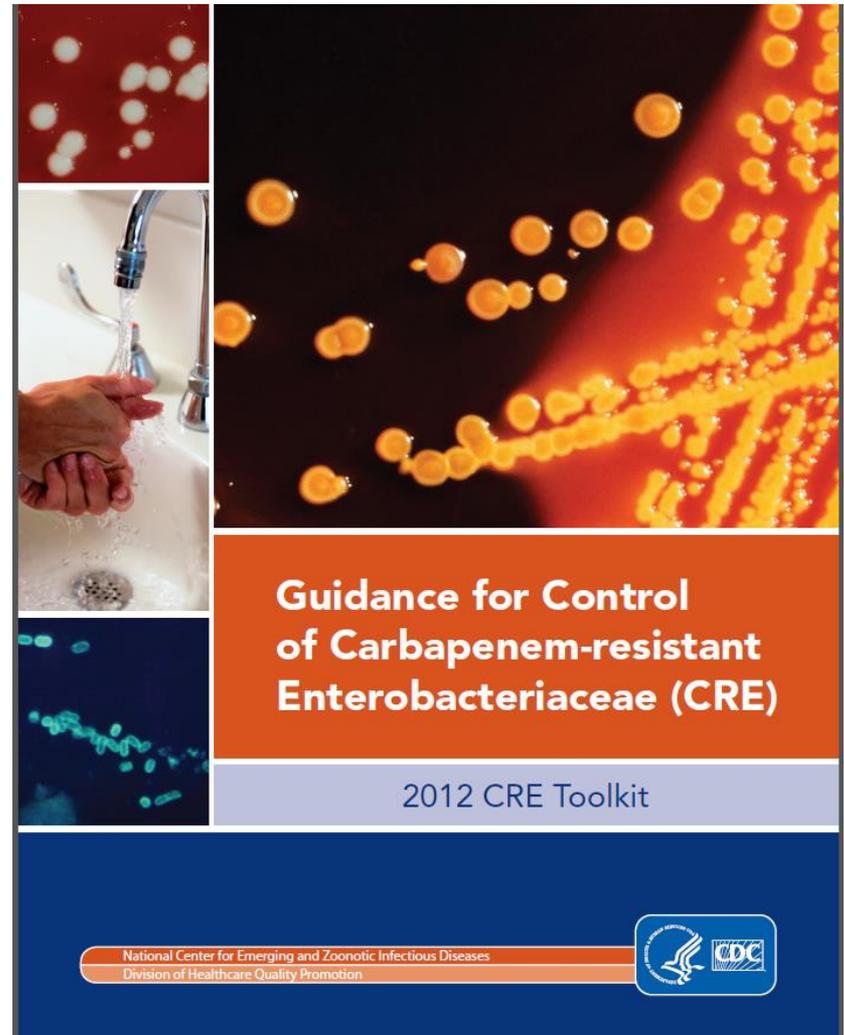
# Definitions of CRE Regional Prevalence

- **No/Rare CRE Identified**
  - Healthcare facilities in region have not identified CRE colonized or infected patients; or some facilities with rare CRE
- **Few CRE**
  - Most healthcare facilities in region do not regularly admit patients with CRE
  - Several facilities in region identify CRE colonized or infected patients on an infrequent basis (e.g. monthly basis or greater)
  - Some facilities with several CRE colonized or infected patients, but surrounded by facilities with only a few or none
- **Common CRE**
  - Majority of healthcare facilities in region have identified cases, and regularly have CRE colonized or infected patients admitted (e.g. CRE detected at least weekly)

# Preventing CRE Transmission

## CDC Toolkit, 2012

- Recommends Core prevention strategies for the control of CRE in ALL hospitals and LTC facilities
- Recommends public health action based on regional CRE prevalence



# CDC Prevention Strategies

## Core Strategies

High levels of scientific evidence

Demonstrated feasibility

- Should become standard practice

## Supplemental Strategies

Some scientific evidence

Variable levels of feasibility

- Consider implementing in addition to Core when infections persist or rates are high

# **CORE** CRE Prevention Strategies for ALL Acute and LTC Facilities

1. Hand Hygiene
2. Contact Precautions
3. Healthcare Personnel Education
4. Minimize Device Use
5. Patient and Staff Cohorting
6. Laboratory Notification
7. Antimicrobial Stewardship
8. CRE Screening

# **SUPPLEMENTAL** Prevention Strategies for Healthcare Facilities with CRE Transmission

1. Active Surveillance Testing
2. Chlorhexidine Bathing

# 1. Hand Hygiene

## CORE

- Ensure healthcare personnel are familiar with proper hand-hygiene and its rationale
  - Make hand hygiene accessible, easy, and convenient
  - Emphasize use of hand hygiene in view of the patient and always prior to patient contact
- Perform adherence monitoring for hand hygiene!
  - Periodically record adherence via direct observation studies
  - Consider routine monitoring via volume of alcohol-based hand rub used
  - When outbreaks occur, assess adequacy of existing policies, procedures, and adherence monitoring practices

## 2. Contact Precautions

**CORE**

- Patients infected or colonized with CRE must be placed Contact Precautions
  - Educate HCP about use and rationale behind Contact Precautions for CRE
  - Perform adherence monitoring for Contact Precautions
- Systems should be in place to identify patients at readmission
- Consider pre-emptive Contact Precautions of patients transferred from CRE high-risk settings or regions
- Notify receiving healthcare facility when transferring patients

# Contact Precautions in Skilled Nursing or Long Term Care Facilities

**CORE**

- Implement contact precautions for LTC residents who are at higher risk for transmitting CRE
  - Totally dependent upon HCP for their activities of daily living
  - Ventilator-dependent
  - Incontinent of stool
  - Wounds with difficult-to-control drainage
- For other CRE colonized residents, the requirement for Contact Precautions might be relaxed
- As with all care, Standard Precautions should be maintained

# Maintain Contact Precautions

## CORE

- The duration of CRE carriage is very extended in comparison to other healthcare pathogens
- An Israeli study (2010) found that among 97 patients positive for CRKP, time to 1<sup>st</sup> negative negative (without subsequent positive)

Mean 387 days and Median 295 days

- Recommendation to discontinue precautions only if 2 screening cultures are both negative

### 3. Healthcare Provider Education

**CORE**

- Education is key to help staff identify those at high risk for CRE acquisition and transmission
  - Review CRE risk factors
  - Describe lack of treatment options
  - Explain necessity for extended contact precautions due to duration of carriage
- AHRQ Toolkit - <http://www.ahrq.gov/cretoolkit>
  - Provides tools for assessment of HCP attitudes and knowledge of CRE control and prevention

## 4. Minimize Device Use

**CORE**

- Presence of medical devices such as catheters, tubes, drains, and ventilators are significant risk factors for CRE
  - Monitor for necessity on a daily basis
  - Require documentation of daily necessity
  - Remove as soon as possible
- Refer to CDC published guidelines for preventing device-associated infections

[www.cdc.gov/hicpac/pdf/CAUTI/CAUTIguideline2009final.pdf](http://www.cdc.gov/hicpac/pdf/CAUTI/CAUTIguideline2009final.pdf)

[www.cdc.gov/hicpac/BSI/BSI-guidelines-2011.html](http://www.cdc.gov/hicpac/BSI/BSI-guidelines-2011.html)

[www.cdc.gov/mmwr/preview/mmwrhtml/rr5303a1.htm](http://www.cdc.gov/mmwr/preview/mmwrhtml/rr5303a1.htm)

## 5. Patient and Staff Cohorting

**CORE**

- CRE patients, whether colonized or infected, should be placed in single rooms when possible
- Implement staff cohorting, dedicating staff to care only for CRE patients during their shift
- If single rooms are not available for all CRE patients:
  - Preference for private rooms should be given to patients at highest risk for transmission such as patients with incontinence, medical devices, or wounds with uncontrolled drainage
  - If room sharing necessary, try to place patients together only if infected with same pathogen / carbapenemase

## 6. Laboratory Notification

**CORE**

- Hospital personnel should receive timely notification from laboratory when a CRE result is identified
  - Preliminary alerts should be sent even before final results are confirmed
  - All final notifications should be received within 24 hours of the initial result

## 7. Antimicrobial Stewardship

**CORE**

- Since 2008, California law requires every acute care hospital to implement a program to monitor “the judicious use of antibiotics”
- Critically important to prevent the spread of CRE
  - Ensure antibiotic prescribed is appropriate for indication
  - Select narrowest spectrum possible
- For assistance, the CDPH HAI Program recently launched “Spotlight on ASPs”
  - Provides criteria for development of basic, intermediate, and advanced tiers of ASPs
  - Lists hospital experts willing to mentor others

[www.cdph.ca.gov/programs/hai/Pages/AntimicrobialStewardshipProgramInitiative.aspx](http://www.cdph.ca.gov/programs/hai/Pages/AntimicrobialStewardshipProgramInitiative.aspx)

## 8. CRE Screening

**CORE**

- Conduct surveillance cultures of patients with epidemiologic links to known CRE patients
- Rationale: Only a minority of patients colonized with CRE will have positive clinical cultures
  - Israeli hospitals found only 5 of 16 patients had positive CRE clinical cultures
  - NY hospital found 2/3 more CRE patients by screening than were identified by clinical culture, resulting in 1400 days of unprotected exposure
- For guidance on performing CRE rectal or peri-rectal swabs, refer to the CDC laboratory protocol at

[www.cdc.gov/ncidod/dhqp/pdf/ar/Klebsiella\\_or\\_E.coli.pdf](http://www.cdc.gov/ncidod/dhqp/pdf/ar/Klebsiella_or_E.coli.pdf)



# 1. Active Surveillance Testing

## SUPPLEMENTAL

- Consider Active Surveillance Testing for:
  - All patients admitted to facility
  - Patients admitted to high-risk settings (e.g. ICU)
  - Patients at highest risk of being colonized with CRE
    - Extended ICU, LTAC hospital or LTC facility stay
    - Extensive antimicrobial exposure
    - Presence of indwelling medical device

## 2. Chlorhexidine Bathing

## SUPPLEMENTAL

- Has been used successfully to prevent some types of HAIs (i.e. CLABSI, CDI) and to decrease colonization with specific MDRO (i.e. MRSA, VRE)
- Consider daily bathing of high risk patients
  - Use either diluted 2% liquid chlorhexidine or chlorhexidine-impregnated wipes
- Usually applied to all patients regardless of CRE colonization status if targeting a specific patient population, e.g. ICU patients

# California Statewide CRE Prevalence Survey



# California CRE Prevalence Survey

## Objectives

1. To educate California hospital infection prevention personnel about CRE
  - Facilitate communication and collaboration between infection prevention and microbiology
2. Determine regional prevalence of CRE in California among general acute care hospitals in 2012
  - Assist local public health and healthcare facilities to better utilize the CDC CRE toolkit

# Methods

- Developed in conjunction with CDC
- All 387 eligible California acute care hospitals including long-term acute care (LTAC) contacted
- Conducted over the phone
  - Because survey data gathered from multiple sources, it took several weeks from initial contact to completion
  - Approximately 15 minutes to complete
- 5 CDPH staff members and 1 volunteer conducted surveys from May 2013 – March 2014

## Methods - continued

Survey included:

- Assessment of CRE infection prevention measures, screening practices, laboratory protocols, and staff awareness
- Hospital prevalence of specific CRE organisms in 2012
  - Definition of CRE: any *Enterobacteriaceae* that tested non-susceptible to a carbapenem
  - Total numbers of *Klebsiella* spp. and *Escherichia coli* isolates
- Collection of 2012 antibiograms
  - Aggregated antimicrobial susceptibility data

# High Variability of CRE *Klebsiella* Prevalence Across California

Hospital Type	Hospitals	Non Susceptible Isolates	Total Isolates	Pooled Mean Prevalence	Percentile Distribution				
					10 <sup>th</sup>	25 <sup>th</sup>	50 <sup>th</sup>	75 <sup>th</sup>	90 <sup>th</sup>
General Acute Care	297	2,264	72,387	3.1%	0%	0%	0%	3%	8%
Long Term Acute Care	22	1,157	2,220	52.1%	2%	25%	41%	66%	76%

Important to note: Even though more than half of all hospitals reported zero resistant isolates in 2012, CRE prevalence varied widely *within regions* and across CA

# Carbapenem Resistant *Klebsiella* species in General Acute Care Hospitals

## General Acute Care Hospitals

	Hospitals Reporting	Non-Susceptible Isolates / Total	Crude CRE Rate*	Adjusted CRE Rate**	95% CI
Los Angeles County	65	1,411 / 17,130	82.4	<b>79.4</b>	70.2-89.2
Rest of CA	223	836 / 53,158	15.7	<b>15.1</b>	13.8-16.4
<b>Total</b>	288	2,264 / 72,489	31.2	<b>32.3</b>	29.6-35.1



\* Per 1000 isolates

\*\* Adjusted for patient days

# Carbapenem Resistant *Klebsiella* species in Long Term Acute Care Hospitals

## Long Term Acute Care Hospitals

	Hospitals Reporting	Non-Susceptible Isolates / Total	Crude CRE Rate*	Adjusted CRE Rate**	95% CI
Los Angeles County	9	966 / 1,373	703.6	<b>527.2</b>	418.8-647.9
Rest of CA	13	191 / 847	225.5	<b>213.0</b>	166.4-265.3
Total	22	1,157 / 2,220	521.2	<b>273.4</b>	229.7-321.0

# Carbapenem Resistant Klebsiella species in Southern California

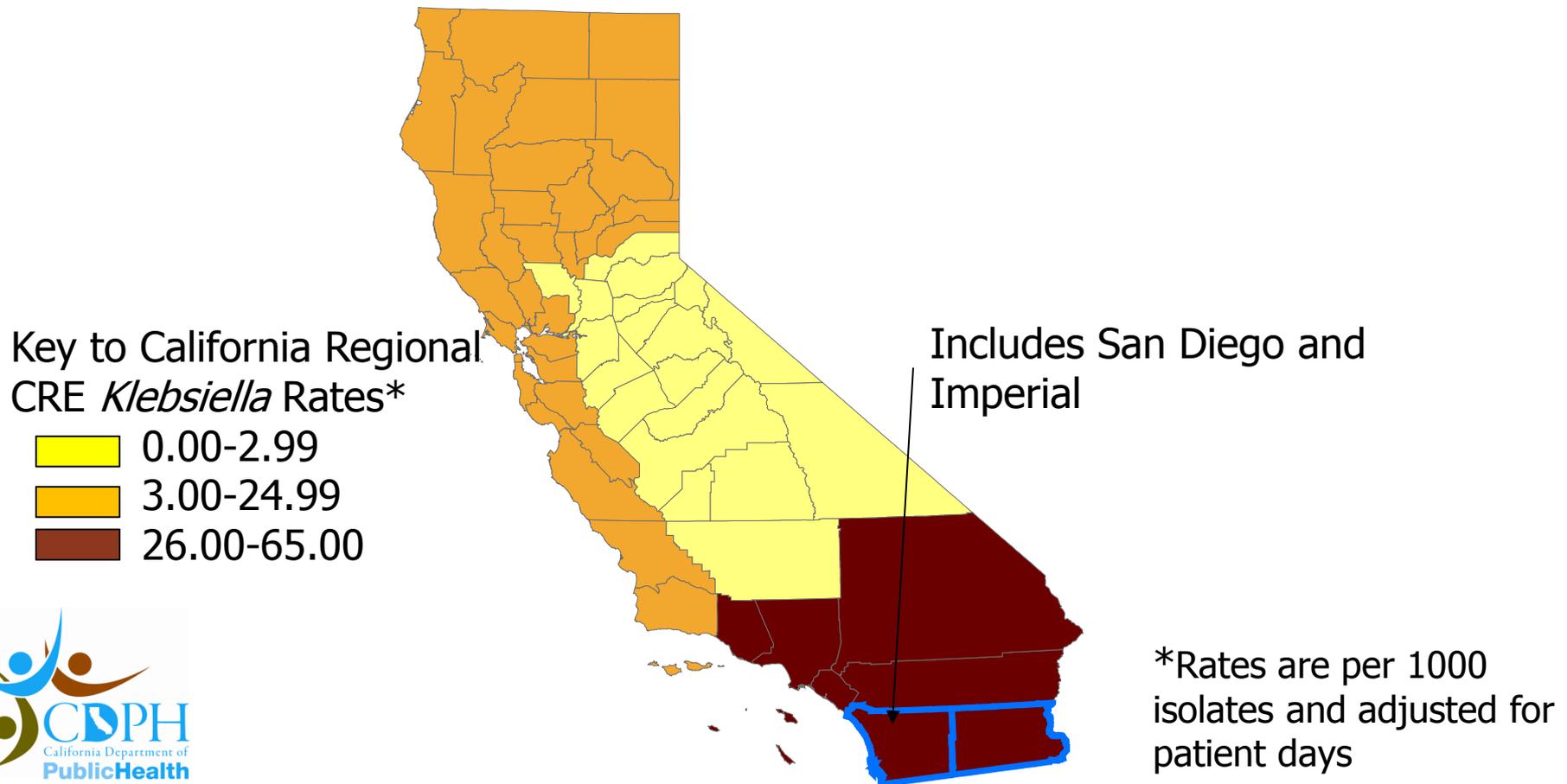
	Hospitals Reporting	Non-susceptible Isolates / Total	Adjusted Prevalence Rate* (95% CI)
<b>Regions</b>			
San Diego Area	18	230 / 8,122	26.4 (22.3-30.7)
Inland Empire	31	270 / 7,472	35.2 (30.5-40.2)
Los Angeles Area	100	1,668 / 25,828	64.6 (57.8-71.8)
<b>Total</b>	<b>149</b>	<b>2,168 / 41,422</b>	

\* All rates are per 1000 isolates tested and adjusted for patient days; LTAC hospitals are excluded; confidence intervals for directly standardized rates calculated using the gamma distribution (Fay and Feuer, 1997)

# San Diego Area

Regional Prevalence of CRE *Klebsiella* Species  
per 1000 Isolates (95% Confidence Interval)

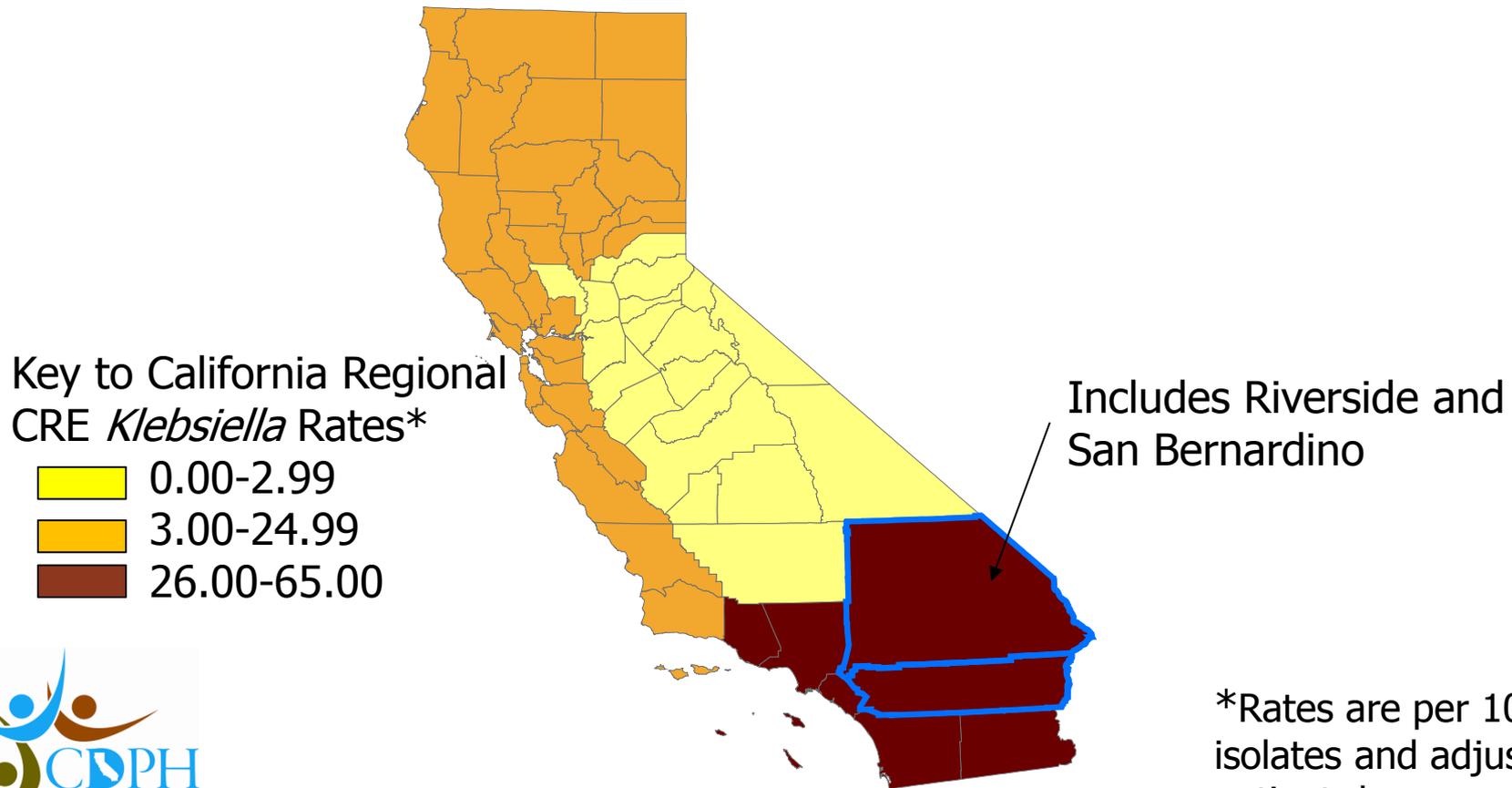
**26.4 (22.3- 30.7)**



# Inland Empire

Regional Prevalence of CRE *Klebsiella* Species  
per 1000 Isolates (95% Confidence Interval)

**35.2 (30.5- 40.2)**



\*Rates are per 1000  
isolates and adjusted for  
patient days

# Los Angeles Area

Regional Prevalence of CRE *Klebsiella* Species  
per 1000 Isolates (95% Confidence Interval)

**64.6 (57.8- 71.8)**



# Preventing CRE Transmission

# CDC CRE Guidance for **Healthcare Facilities** When *First* CRE Patient Identified

Step-by-step process to follow when new CRE-colonized or CRE-infected patient is identified

## **1. Notification**

- Notify appropriate personnel, including both clinical and infection prevention staff
- Notify public health if indicated

## **2. On the Unit/Ward**

- Place patient in Contact Precautions in single room
- Reinforce hand hygiene and use of Contact Precautions on affected ward /unit
- Educate healthcare personnel staff about preventing CRE transmission

### 3. Screening

- Test epidemiologically linked patients (i.e. roommates) using stool or peri-rectal cultures
- Consider point prevalence survey of affected unit
- Consider pre-emptive precautions for epi-linked patients while awaiting test results

### 4. Evidence of Transmission

- If screening cultures identify additional CRE colonized or infected patients, consider additional surveillance cultures of contacts or point prevalence surveys
- Implement cohorting CRE patients and dedicated staffing

### 5. Patient Transfer

- Confirm that Contact Precautions are continued on new unit
- Ensure CRE status shared if patient transferred to another facility



# CDC CRE Guidance for **State/Local Public Health Departments**

## Regions With **No** CRE Identified Emphasis on regional surveillance and education.

### I. **Perform Regional Surveillance and Provide Feedback**

- A. Consider making CRE reportable - OR - Survey healthcare facilities by phone or email
- B. If **NO** CRE cases are identified in region:
  - Feedback results to healthcare facility IPs and lab directors
  - Promote facility implementation of CRE prevention strategies

If CRE cases **ARE** identified in region:

  - Implement appropriate regional strategy depending if CRE "few" or "common"
- C. Repeat survey at least quarterly if CRE in neighboring jurisdictions. Otherwise, repeat at least every 6 months.

### II. **Educate ALL Healthcare Facilities**

- Explain importance of CRE and provide updates on national and/or neighboring regional prevalence and epidemiology
- Review recommended surveillance and prevention measures
- Increase vigilance for CRE detection

## Regions With **FEW** CRE Identified

**Regions where cases remain uncommon. Emphasis on preventing further transmission and widespread emergence. Target select facilities.**

### I. **Perform Regional Surveillance and Provide Feedback**

A. CRE confirmed by survey or reports

- B. Feedback results to healthcare facility IPs, lab directors, and facility administrators by email or letter
- Consider publication of results by facility name, area, type
  - Engage state hospital association, QIO, and other prevention partners to facilitate communication with facility leaders
  - Promote facility implementation of CRE prevention strategies

C. Repeat CRE surveillance and feedback quarterly

### II. **Educate ALL Healthcare Facilities**

- Explain importance of CRE and provide updates on national and/or neighboring regional prevalence and epidemiology
- Review recommended surveillance and prevention measures
- Increase vigilance for CRE detection

### III. **Inter-facility Communication**

Ensure facilities complete an Inter-facility Transfer Form when transferring CRE patients

## Regions With **FEW** CRE Identified – *continued*

### IV. Infection Prevention

For facilities **WITH** CRE:

1. Engage facility administrators to prioritize CRE prevention
2. Review practices to ensure core CRE prevention measures are in place
3. Provide in-service training
4. Ensure CRE screening is being performed
5. If CRE rates do not decrease, consult CDC for more guidance

For facilities **WITHOUT** known CRE but located in region where CRE are present:

1. Engage facility administrators, ensure control plan, and reinforce CRE Core prevention strategies
2. Guide implementation of CRE screening and preemptive Contact Precautions for patients admitted from
  - Facilities with ongoing CRE transmission
  - LTAC hospitals or with CRE risk factor

## Regions Where CRE are **COMMON**

Emphasis on implementation of core and supplemental prevention measures across **ALL** acute care and LTC facilities.

### I. Dedicated Personnel to Engage HC Facilities

- Assign specific personnel to this task
- Form advisory panel if additional technical support is needed
- Engage all facility administrators and IP personnel early in process
  - Hospital Association, QIOs, and other relevant partners

### II. Perform Regional Surveillance and Provide Feedback

- A. Perform steps for CRE surveillance and feedback of results to IP and/or lab directors and facility administrators
- B. Determine if certain CRE events should be made reportable:
- C. Repeat survey at least quarterly

### III. Inter-facility Communication

Ensure facilities complete an Inter-facility Transfer Form when transferring CRE patients

## Regions With **COMMON** CRE Identified – *continued*

### IV. Infection Prevention

- A. Reinforce Core prevention measures in all facilities
  - Work closely with IPs to review practices
  - Provide in-service training
- B. Consider Supplemental measures in all facilities
  - Active surveillance testing and preemptive Contact Precautions for patients admitted
    - from facilities with ongoing CRE transmission or high CRE prevalence
    - from LTAC hospitals or with CRE risk factors
    - to high-risk units (ICUs)
  - Chlorhexidine bathing on high-risk patients
- C. Assess Compliance to Prevention Measures (monthly)
  - Share performance measures with facility
  - Provide in-service training, as needed

### V. Assess Efficacy of Infection Prevention Measures

- If CRE rates do not decrease
  - Intensify efforts in select facilities as needed
  - Implement additional interventions in consultation with CDC or state/regional experts

# Effective CRE Prevention Will Require Local Partnerships

- Regional prevention collaboratives should include general acute care and LTAC hospitals, SNF/LTC facilities, and public health

*“An effective intervention at containing the spread of CRE should ideally be implemented before CRE have entered a region, or at the very least, immediately after its recognition.*

*Policy makers and public health authorities must ensure the early recognition and coordinated control of CRE.”*



# So How are California Hospitals Doing with CRE Prevention Currently?

# Survey Results: CRE Awareness and Surveys

<b>Awareness and Surveys</b>	<b>Adherent / Total Responses</b>	<b>%</b>
Read CDC CRE Health Alert Network (HAN)	281 / 315	89
Uses inter-facility transfer form	236 / 321	74
Conducted point prevalence survey	12 / 326	4
Found any CRE	2 / 11	18
Used CDC protocol	7 / 11	64

# Survey Results: CRE Screening Practices

Screening Practices	Adherent/ Total Responses	%
Ask about recent travel history	128 / 324	39
Ask about healthcare exposures in past 6 months	100 / 323	31
Epidemiologically linked to a positive CRE patient	117 / 303	39
Screen for CRE upon Admission	23 / 325	7
All patients	1 / 325	<1
Patients hospitalized overnight outside US <6 mo	4 / 325	1
Patients admitted from LTAC hospital	5 / 325	2
Patients admitted from SNF/LTC facility	5 / 325	2
Patient with history of MDRO	1 / 325	<1
Patient admitted from ICU	3 / 325	1

# Survey Results: Laboratory

<b>Laboratory Notification and Record Review</b>	<b>Adherent / Total Responses</b>	<b>%</b>
Timely Notification	298 / 326	91
Preliminary Alerts	203 / 298	68
Estimated Time till Notification		
<24 hours	224 / 292	77
24-48 hours	53 / 292	18
>48 hours	8 / 292	3
Retain Isolates	156 / 322	48
Perform Micro Record Review	122 / 326	37
Found any CRE	31 / 122	25

# Lab Capacity

**Save all CRE lab isolates!**

- Either local public health or CDPH may provide additional reference testing
- CDPH is developing a roadmap and increasing personnel to enhance lab capacity at the state level

# CRE Outbreaks

- Suspected CRE outbreaks or clusters (i.e. an unusual occurrence of CRE) must be reported to local public health
- CRE outbreaks have been serious and difficult to control
  - Outbreak of CRE *Klebsiella* at National Institutes of Health hospital in 2011 resulted in person-to-person transmission causing 18 infections and 11 deaths
  - Outbreak of CRE *E. coli* in Chicago area in 2013 associated with a single endoscope, resulted in 8 infections and 44 colonizations

# CA Regional Prevalence of CRE *Klebsiella*

	Number of Hospitals	No. Hosp with $\geq 1$ CRE Isolate	Nonsusceptible Isolates	Total Isolates	Adjusted Prevalence Rate* (95% CI)
<b>Regions</b>					
Sierras	5	0	0	467	0.00 (0.0-0.1)
Sacramento Metro	13	2	2	3,643	0.92 (0.1-2.6)
San Joaquin Valley	36	11	27	9,102	2.59 (1.6-3.8)
Northern California	29	6	13	4,244	4.18 (1.8-7.6)
Bay Area	49	18	41	11,596	5.20 (3.4-7.3)
Central Coast	16	4	13	2,015	9.72 (4.8-16.4)
San Diego Area	18	16	230	8,122	26.36 (22.3-30.7)
Inland Empire	31	24	270	7,472	35.20 (30.5-40.2)
Los Angeles-Orange-Ventura	100	75	1,668	25,828	64.65 (57.8-71.8)
<b>Total</b>	297	156	2,264	72,387	34.60 (31.8-37.6)

# In Summary: California CRE Prevalence and Prevention

1. CRE prevalence is significantly higher in the Los Angeles County than other regions of California
2. CRE prevalence is significantly higher in LTAC hospitals than other general acute care hospitals
3. CRE awareness is high among California IPs, however adherence to CDC prevention guidelines varies widely
4. Less than half of respondents are screening other patients when a CRE case is identified
5. Regional efforts to contain CRE will require collaboration between public health and healthcare facilities

**A heartfelt thanks to all of California hospital personnel who participated in the survey and continue the fight against CRE**

For more information, please contact  
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