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[Pollev.com/spice](https://Pollev.com/spice)

Questions will appear on your screen as we go through the presentations.

For your convenience, you may prefer to download the app: Poll Everywhere

# I understand the process for CRE reporting and surveillance



# I can identify resources and key stakeholders related to CRE response



# I can describe containment strategies for CRE

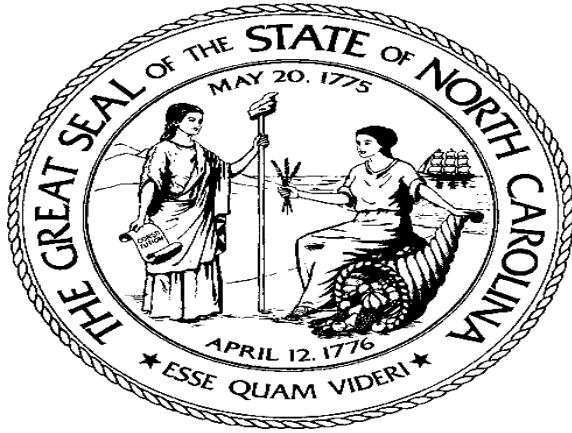


# I am comfortable assessing infection prevention measures to prevent CRE transmission



# I understand the process for initiating CRE colonization screening





# **CRE Surveillance, Identification, Containment & Response**

**NC DPH SHARPPS**

**Communicable Disease Branch**

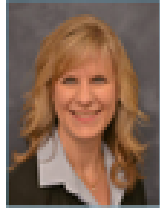
**North Carolina Division of Public Health**

**December 5, 2018**

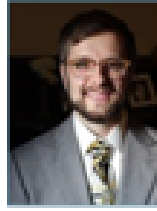


# SHARPPS

## *Surveillance for Healthcare Associated & Resistant Pathogens Patient Safety Program*



**Jennifer  
MacFarquhar**  
Program Director



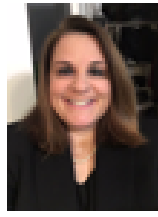
**James  
Lewis**  
Medical Director



**Heather  
Dubendris**  
Epidemiologist



**Katie  
(Steider)  
Palladino**  
Epidemiologist



**Deborah  
Dolan**  
Health Educator,  
Campaigns Coordinator



**Savannah  
Carrico**  
Epidemiologist

**Coming Soon!**  
Epidemiology Program  
Manager





# Objectives

- **Discuss the public health significance of CRE and the process for surveillance and detection**
- **Discuss unified response to CRE**
- **Describe containment strategies for CRE in individual facilities**

# Disclosure

- The presenters for this session have no financial conflicts of interest to disclose

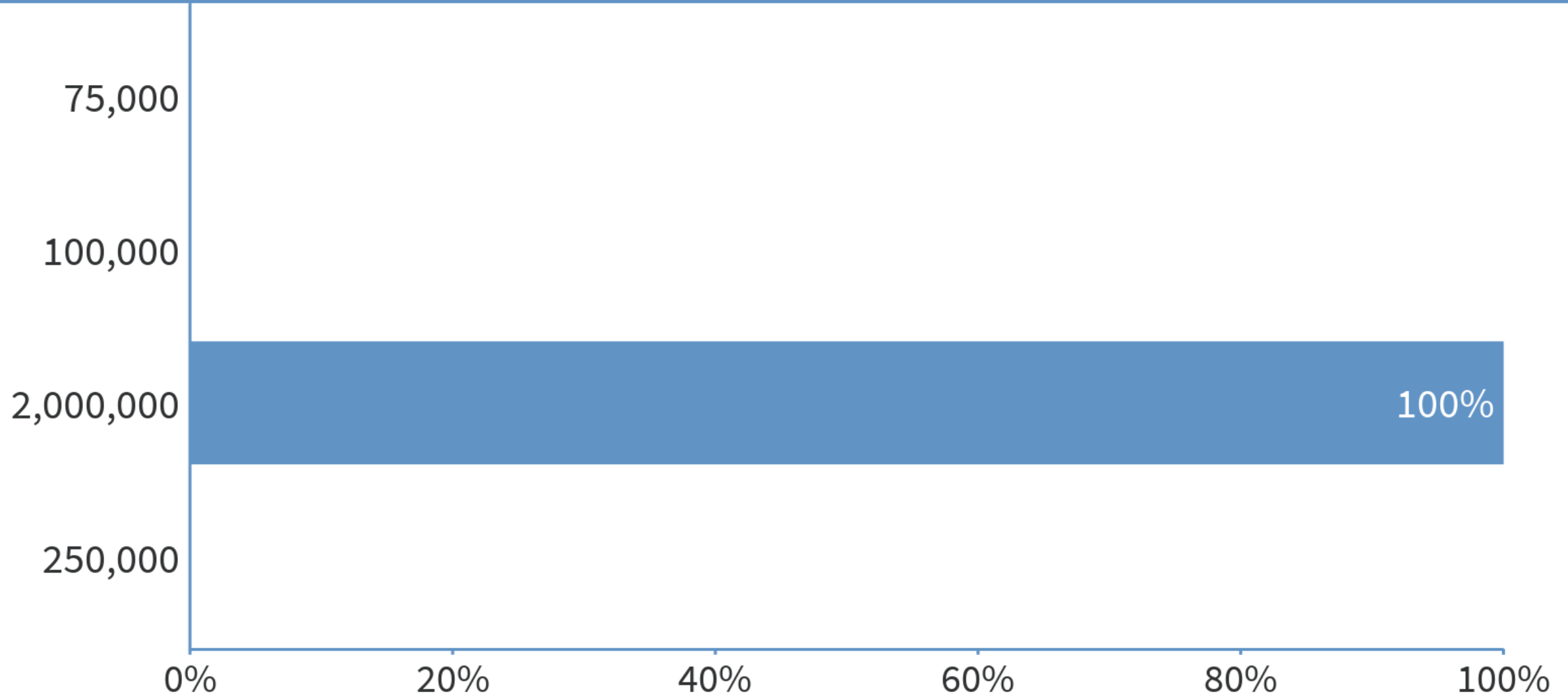


# What is your background and training?

# Multidrug-Resistant Organisms (MDROs)



# According to CDC, what is the estimated number of infections caused by antibiotic resistant pathogens in the U.S. annually?





## MDROs

**MRSA** - Methicillin-resistant  
*Staphylococcus aureus*

**MDR Acinetobacter** –  
Multi-drug resistant  
*Acinetobacter*

**MDR Pseudomonas** –  
Multi-drug resistant  
*Pseudomonas*

**CRE** - Carbapenem-Resistant  
Enterobacteriaceae

**VRE** - Vancomycin-Resistant  
Enterococci

**C Diff** - *Clostridium difficile*

**ESBLs**- Extended Spectrum  
Beta-Lactamase Producers



## MDROs

**MRSA** - Methicillin-resistant  
*Staphylococcus aureus*

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Multi-drug resistant  
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Enterobacteriaceae

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**ESBLs**- Extended Spectrum  
Beta-Lactamase Producers



# Carbapenem-Resistant Enterobacteriaceae (CRE)

- Enterobacteriaceae = gram negative bacteria found in the digestive tract
  - *E. Coli*
  - *Klebsiella spp.*
- CRE = Enterobacteriaceae resistant to carbapenem antibiotics





# Carbapenems

- Class of **Beta-lactam** antibiotics
  - Ertapenem
  - Meropenem
  - Imipenem
  - Doripenem
- Usually reserved to treat drug-resistant infections

BE ANTIBIOTICS AWARE: SMART USE, BEST CARE



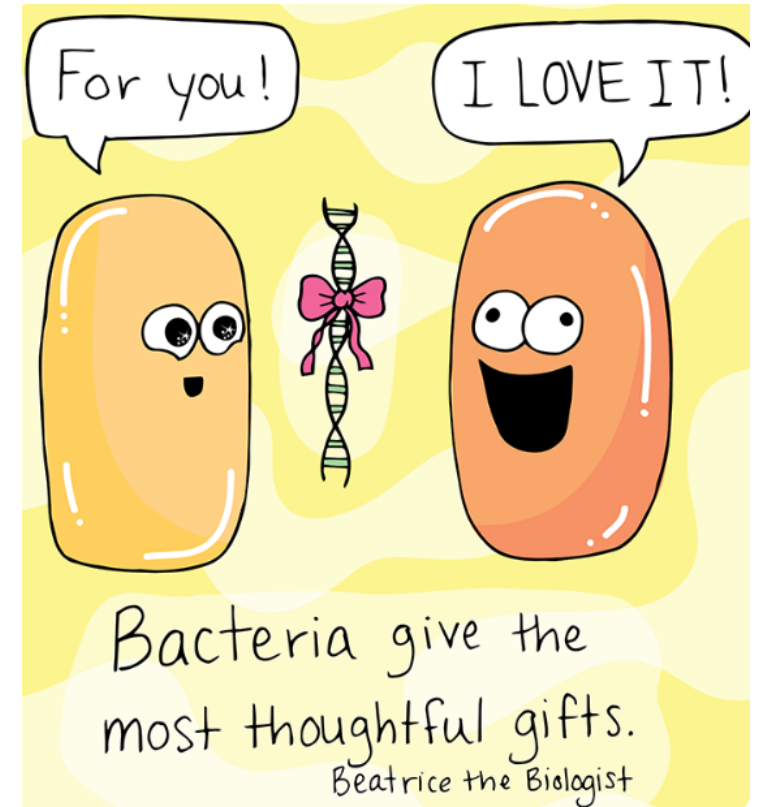
*Artwork submitted by 11th grader, Mallori Mull of Mount Holly, NC, Winner of the 2017 NC Get Smart Artwork Competition*

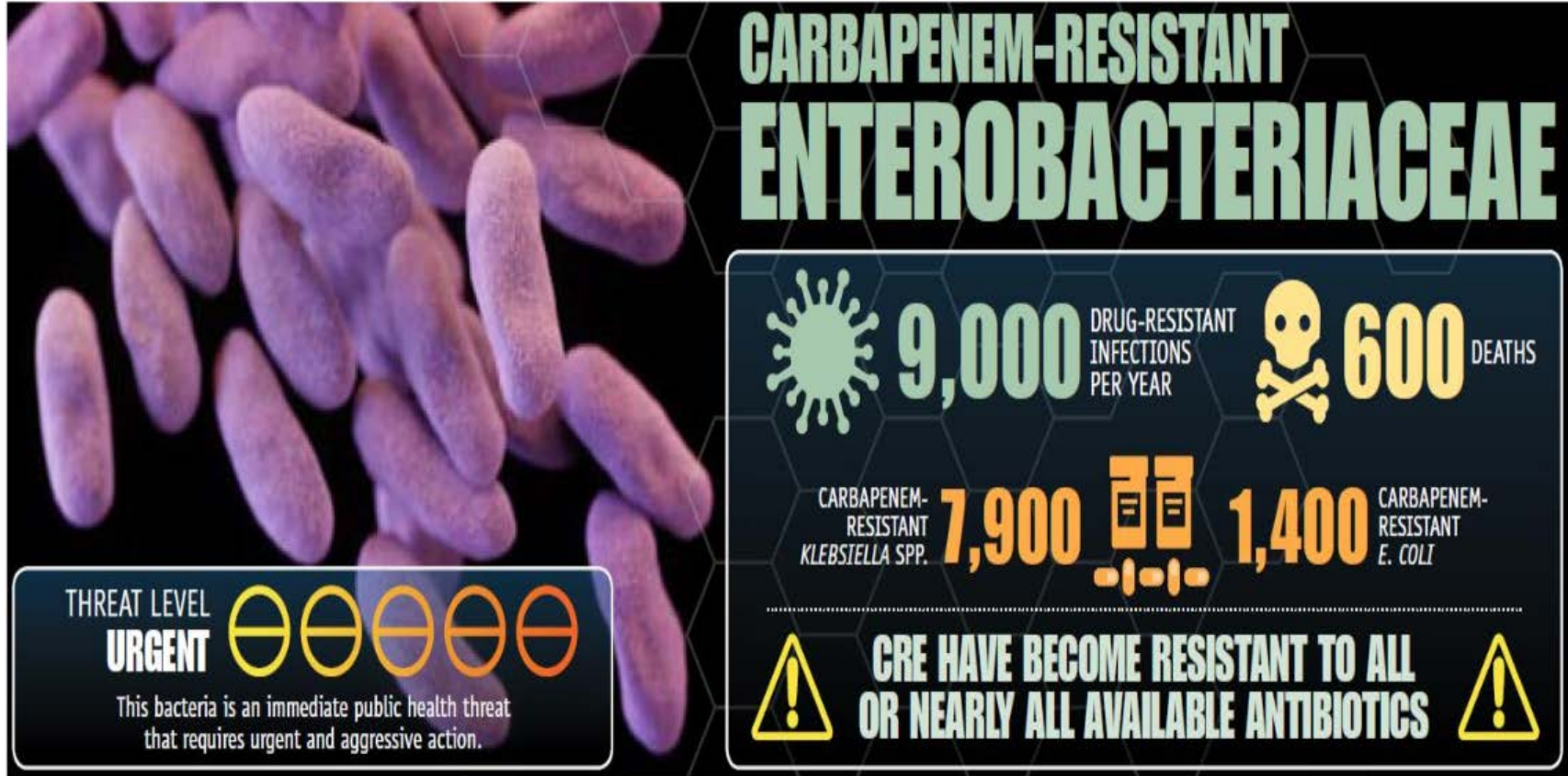
For more information, visit the NC Get Smart Campaign:  
<http://epi.publichealth.nc.gov/cd/antibiotics/campaign.html>



# Carbapenemase producing CRE (CP CRE)

- Carbapenemase = enzyme that can break down carbapenem antibiotics
  - Klebsiella pneumoniae carbapenemase (KPC),
  - New Delhi metallo- $\beta$ -lactamase (NDM),
  - Verona integron encoded metallo- $\beta$ -lactamase (VIM),
  - Imipenemase metallo- $\beta$ -lactamase (IMP)
  - Oxacillinase-48 (OXA-48)
- Mobile resistance elements





CDC: Antibiotic Resistance Threats in the United States, 2013



# What NC DPH is doing:

- **Detect MDROs**
  - Increased awareness
  - Sentinel surveillance
  - Testing at SLPH
  - Colonization screening
- **Ensure rapid response & containment**
  - Systematic response to even single cases
  - Infection prevention assessments
  - Inter-facility communication
  - Screening for colonization
- **Stewardship efforts**
  - Antimicrobial resistance subcommittee
  - Get Smart to Be Antibiotics Aware
  - STAR partners
- **Education**
  - Webinars
  - Toolkits
  - Presentations
  - Guidance documents



# What NC DPH is doing:

- **Detect MDROs**

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- **Ensure rapid response & containment**

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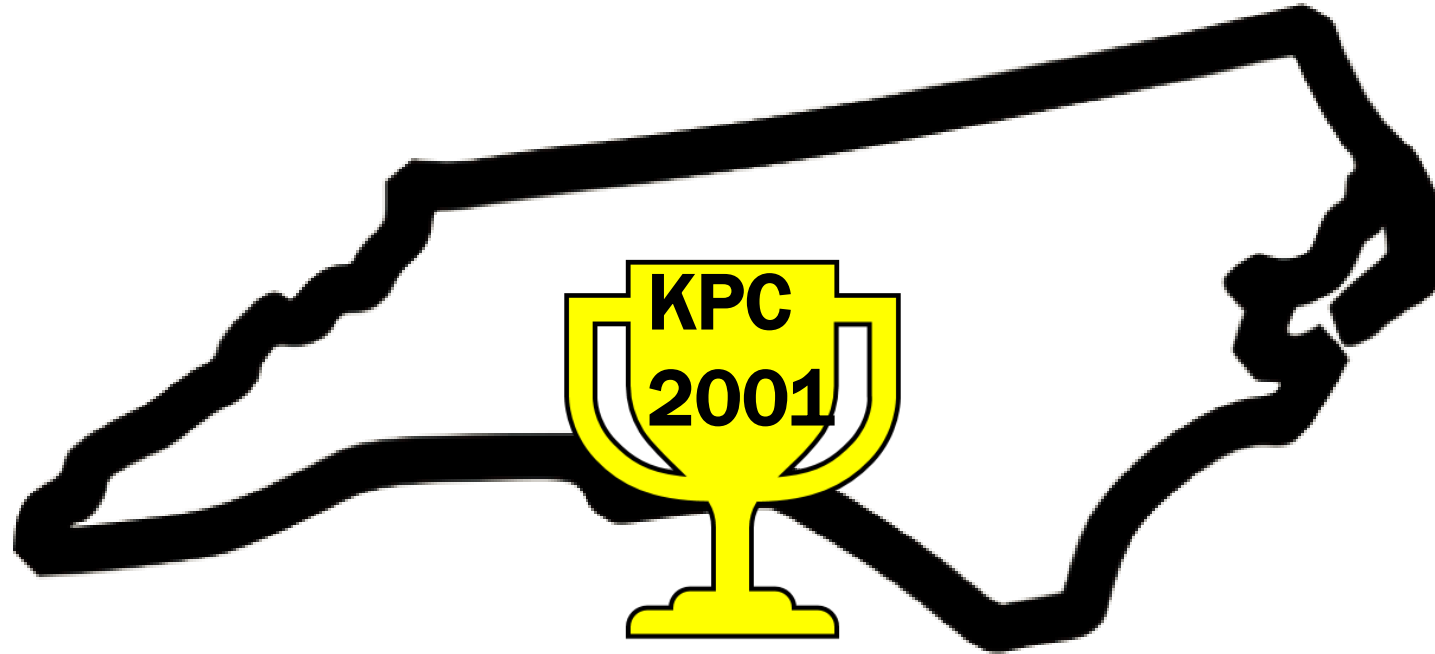
- **Education**

- Webinars
- Toolkits
- Presentations
- Guidance documents

# **DETECTION & SURVEILLANCE**

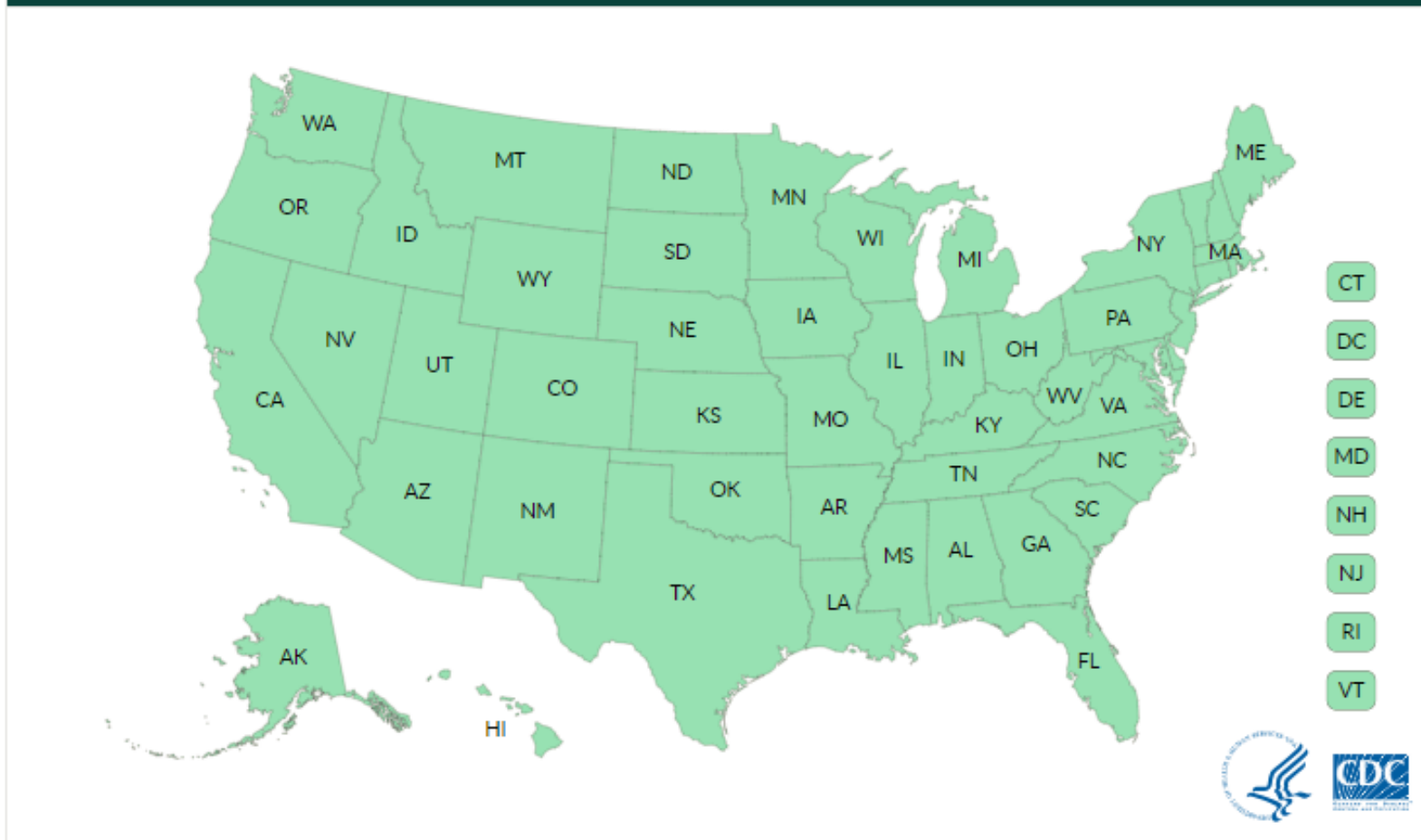
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# Coming in First



# KPC has been reported in all 50 states

Patients with KPC-producing *Carbapenem-resistant Enterobacteriaceae* (CRE) reported to the Centers for Disease Control and Prevention (CDC) as of December 2017, by state



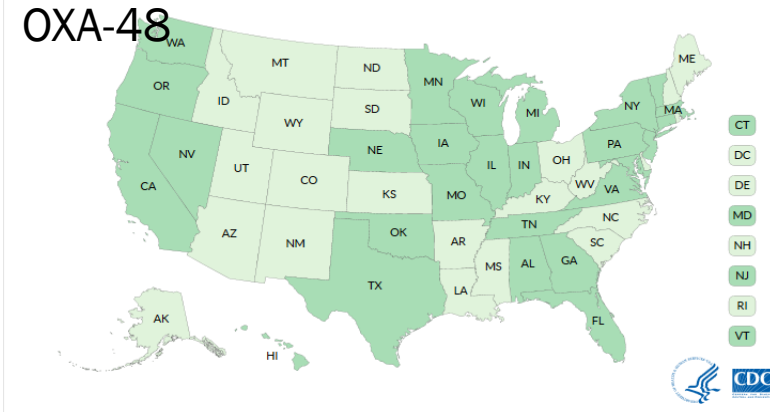
KPC enzyme

- None
- Reported

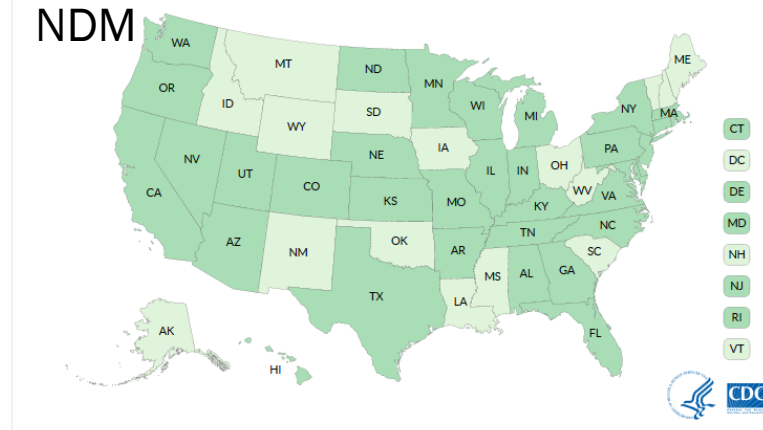


# Detection of other CP-CRE varies by state

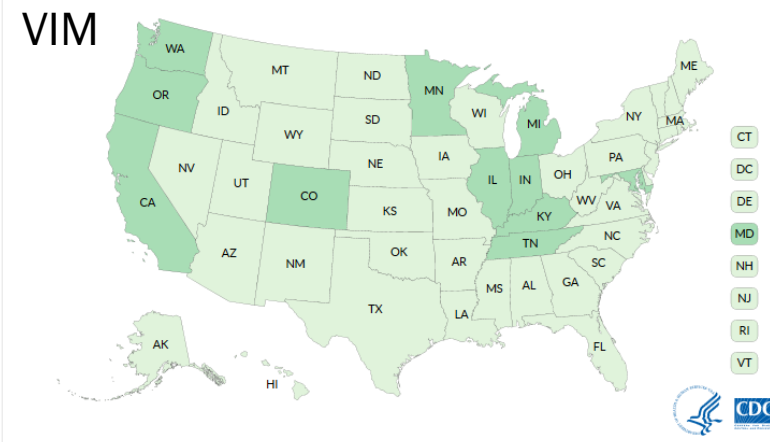
Patients with OXA-48-Type-producing *Carbapenem-resistant Enterobacteriaceae* (CRE) reported to the Centers for Disease Control and Prevention (CDC) as of December 2017, by state



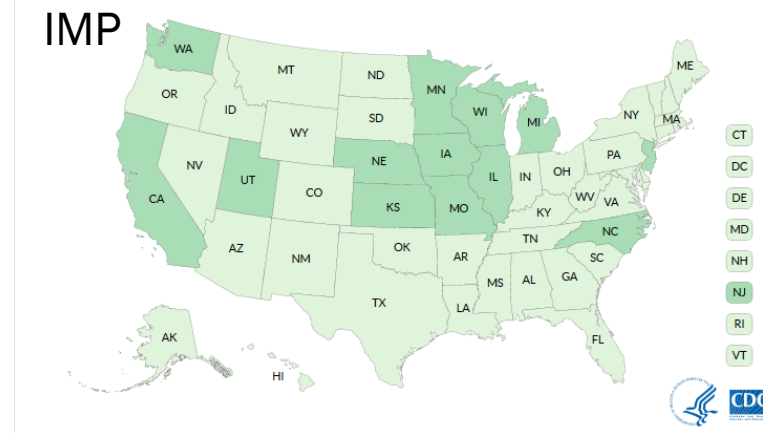
Patients with NDM-producing *Carbapenem-resistant Enterobacteriaceae* (CRE) reported to the Centers for Disease Control and Prevention (CDC) as of December 2017, by state



Patients with VIM-producing *Carbapenem-resistant Enterobacteriaceae* (CRE) reported to the Centers for Disease Control and Prevention (CDC) as of December 2017, by state



Patients with IMP-producing *Carbapenem-resistant Enterobacteriaceae* (CRE) reported to the Centers for Disease Control and Prevention (CDC) as of December 2017, by state



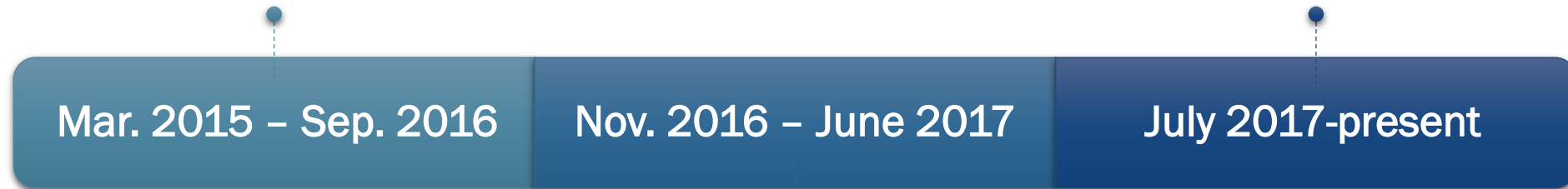
Maps are routinely updated and available from : <https://www.cdc.gov/hai/organisms/cre/trackingcre.html>

# Surveillance in NC



Sentinel site surveillance

Targeted recruitment for sentinel surveillance,  
special projects and outbreak response.



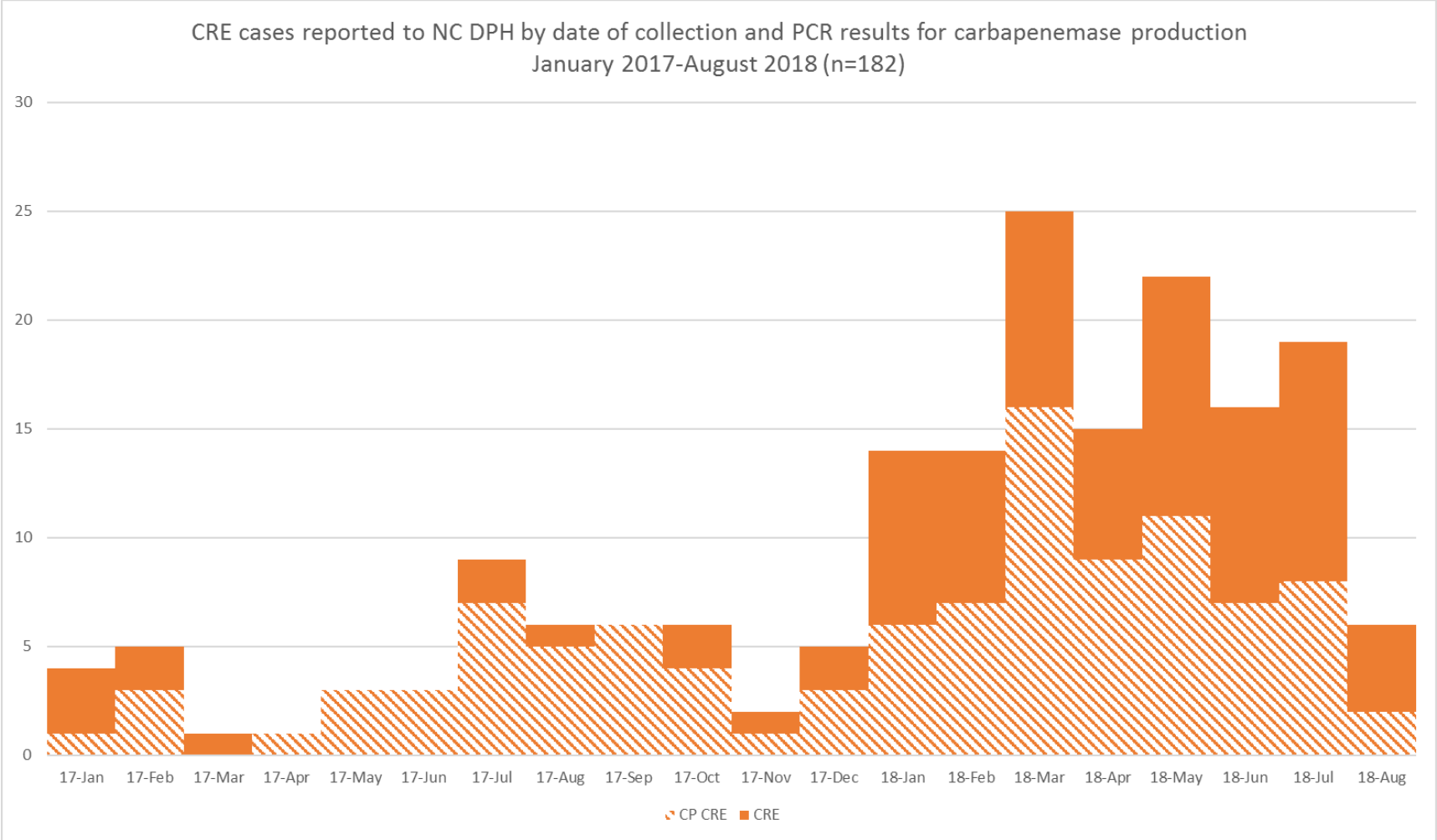
Mar. 2015 – Sep. 2016

Nov. 2016 – June 2017

July 2017-present

Accepted Isolates but did not actively recruit  
sites

# Sentinel surveillance efforts



\*Excludes duplicate CRE (Same Carbapenemase/organism; repeat clinical isolates in a 12 month period; screening results subsequent to a clinical result)



# Sentinel surveillance efforts

**57%** of reported CRE reported to NC DPH are carbapenemase producing

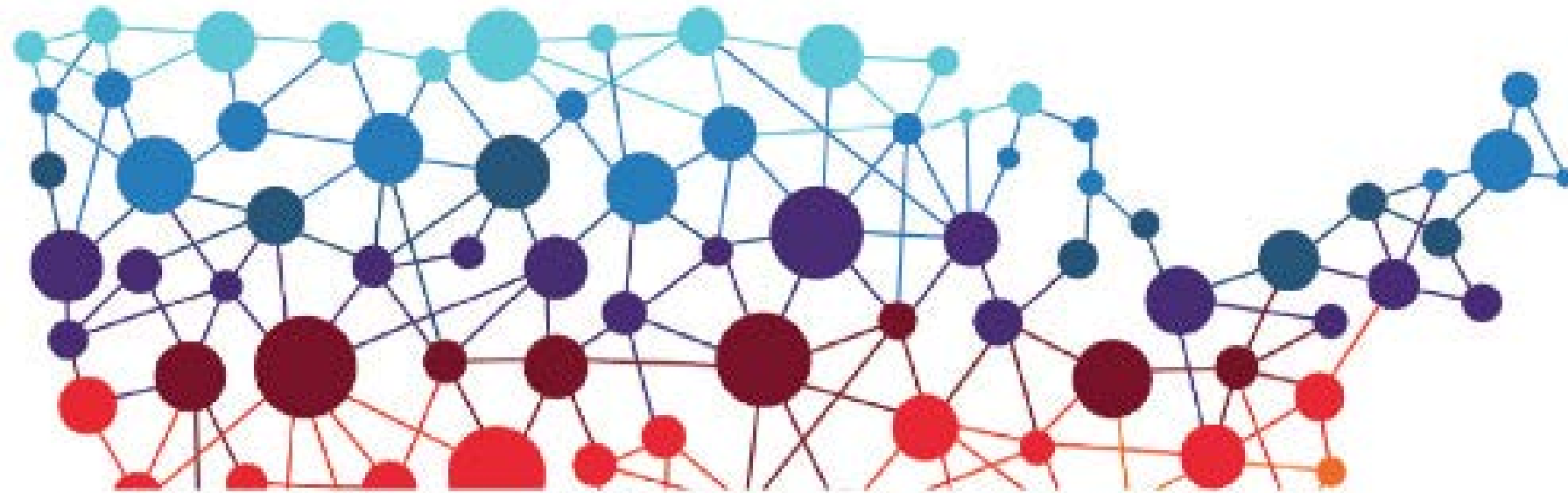
# Sentinel surveillance efforts

The Prevalence of types of carbapenemases among reported CP-CRE, North Carolina January 2017-August 2018  
(n=103)



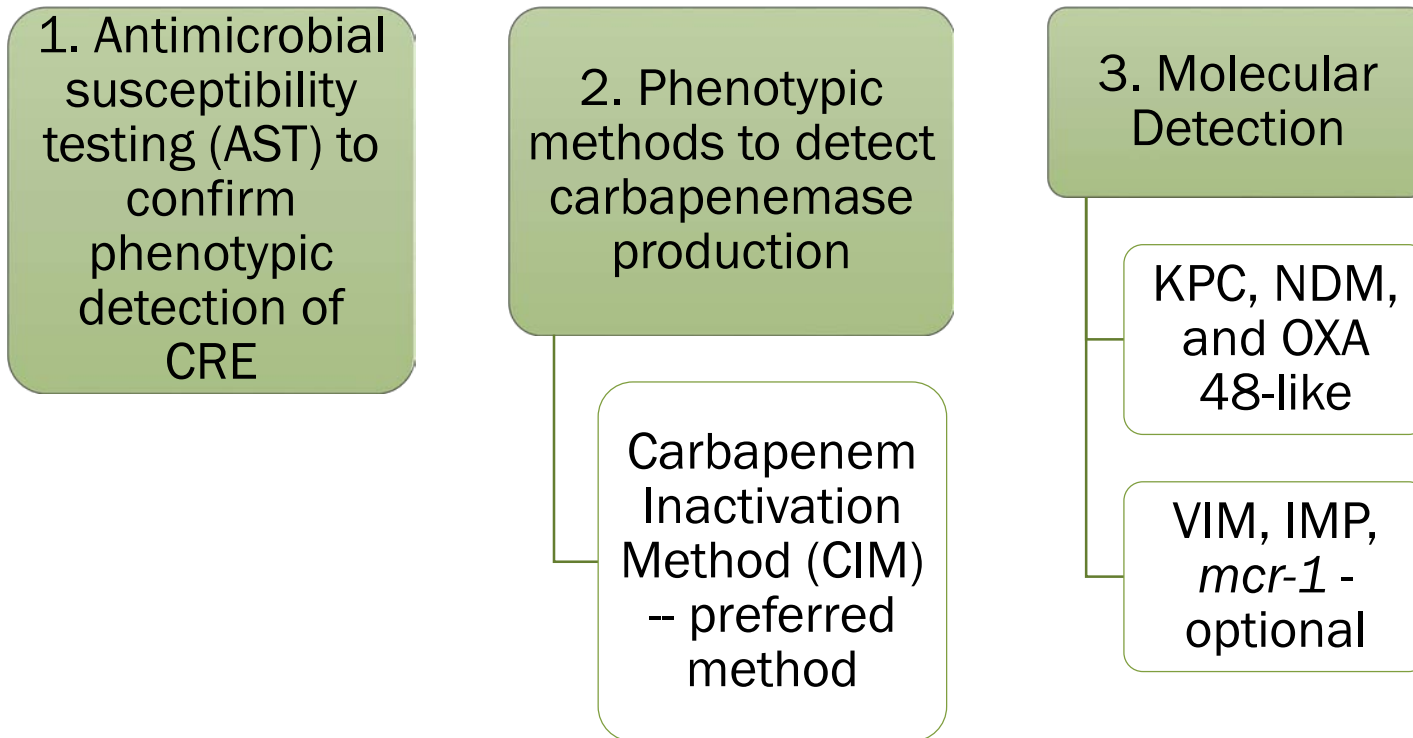


# Utilizing the Antibiotic Resistance Lab Network (ARLN) to detect colonization:



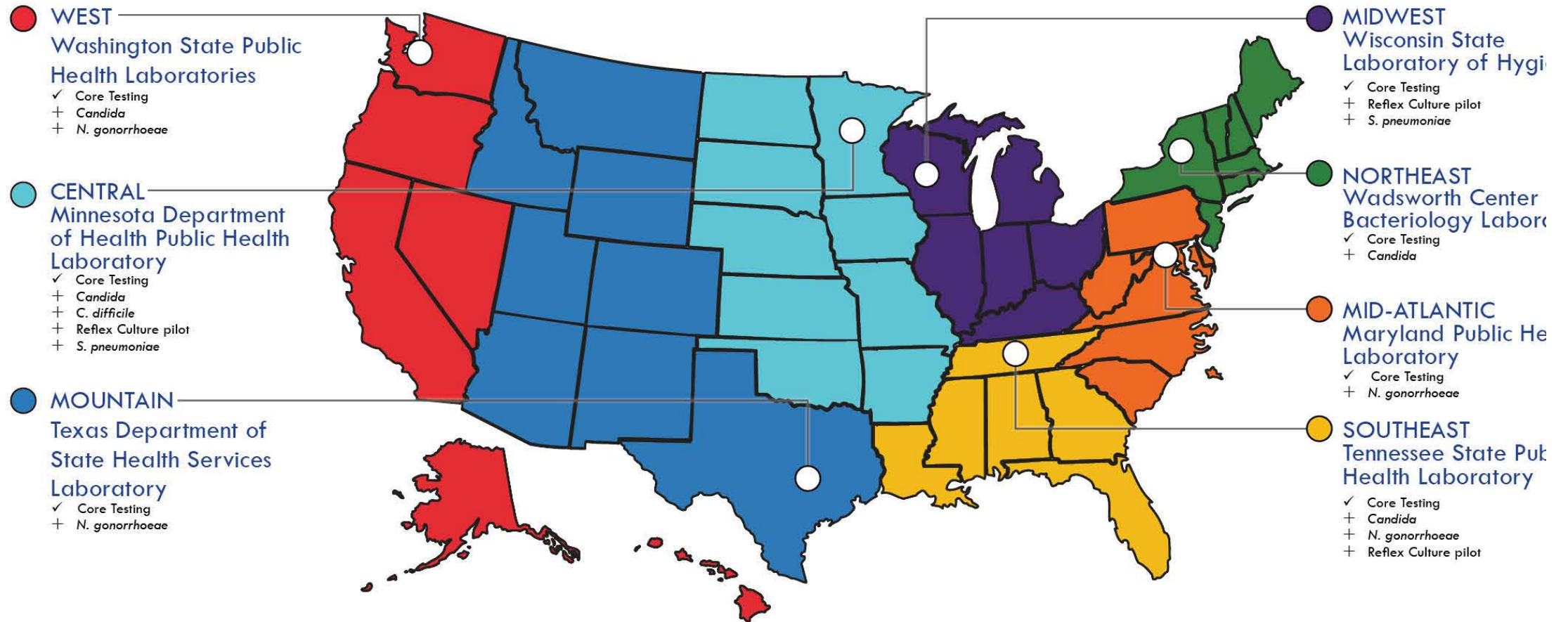
**ARLAB**network

# NC SLPH funded to characterize CRE



# Antibiotic Resistance Lab Network (ARLN)

CDC Antibiotic Resistance Laboratory Network: 7 Regional Labs





# **ADDITIONS TO 10A NCAC 41A .0101**

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**Effective October 1, 2018**

# Additions to 10A NCAC 41A .0101

## Additions include:

- Carbapenem-resistant Enterobacteriaceae (CRE) – 24 hours
- *Candida auris* – 24 hours

## Reporting will:

- Facilitate early detection, rapid response and containment
- Prevent transmission
- Provide data to develop and implement prevention and control measures



# What to report?

- **Identification of CRE from a clinical specimen associated with either infection or colonization –AND –**



# What to report?

- Identification of CRE from a clinical specimen associated with either infection or colonization –AND –
- All susceptibility results (if available) – AND –



# What to report?

- Identification of CRE from a clinical specimen associated with either infection or colonization –AND –
- All susceptibility results (if available) – AND –
- All phenotypic or molecular test results (if conducted and available)



# For the purposes of reporting, Carbapenem-Resistant *Enterobacteriaceae* (CRE) are defined as:

(1) *Enterobacter* spp., *E.coli* or *Klebsiella* spp. positive for a known carbapenemase resistance mechanism or positive on a phenotypic test for carbapenemase production

or

(2) *Enterobacter* spp., *E.coli* or *Klebsiella* spp. resistant to any carbapenem in the absence of carbapenemase resistance mechanism testing or phenotypic testing for carbapenemase production.



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# Lab reports: look for organism identity

- Organism identification
  - May use a culture or “NAAT” (“nucleic acid amplification test”) or “PCR”

<b>Order</b>		<b>CULTURE, URINE [URC] (Order 308868132)</b>	
<b>Ordering Provider</b>			
Authorizing [REDACTED]			
<b>Acknowledgement Info</b>			
For Placing Order	At 05/17/17 1927	Acknowledged By [REDACTED]	Acknowledged On 05/17/17 1957
Task		Completed by [REDACTED]	Date/Time Wed May 17, 2017 7:29 PM
<b>Order Info</b>			
Priority: STAT	Start: 05/17/17 1927	Process Instructions: ** Minimum Specimen Requirements: 25 ML Urine ** **Submit urine in a container with NO preservative** **Use Orange Screw-capped urine cup, White Screw-capped urine tube or Red top tubes**	
<b>Order Frequency</b>			
Antibiotic		Organism <b>&gt;100,000 cfu/ml enterobacter cloacae</b>	Organism
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final
<b>Lab and Collection</b>			
CULTURE, URINE on 6/3/2017			
<b>Result History</b>			
CULTURE, URINE on 6/6/2017			
<b>Reviewed by List</b>			
[REDACTED]			
<b>View SmartLink Info</b>			
Culture, Urine (Order #308868172) on 6/3/17			
Ordering Provider NPI ID			





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# Lab reports: look for susceptibility results

- Antimicrobial susceptibility results
  - Also called “MICs” (“minimum inhibitory concentration”) with “interps” (“interpretation”)

**Order** CULTURE, URINE [URC] (Order 308868132)

---

**Ordering Provider**  
Authorizing [REDACTED]

---

**Acknowledgement Info**

For Placing Order	At 05/17/17 1927	Acknowledged By [REDACTED]	Acknowledged On 05/17/17 1957
Task Unit Sec Ack		Completed by [REDACTED]	Date/Time Wed May 17, 2017 7:29 PM

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Priority: STAT	Start: 05/17/17 1927	Process Instructions: ** Minimum Specimen Requirements: 25 ML Urine ** **Submit urine in a container with NO preservative** **Use Orange Screw-capped urine cup, White Screw-capped urine tube or Red top tubes**
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**Order Frequency**

Antibiotic	Organism	Organism
	>100,000 cfu/ml cloacae	enterobacter
AMP/SULBACTAM	MIC RESISTANT	Final
AMPICILLIN	MIC RESISTANT	Final
AUGMENTIN	MIC RESISTANT	Final
CIPROFLOXACIN	MIC RESISTANT	Final
ERTAPENEM	MIC RESISTANT	Final
GENTAMICIN	MIC SUSCEPTIBLE	Final
NITROFURANTOIN	MIC SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC SUSCEPTIBLE	Final

---

**Lab and Collection**  
CULTURE, URINE on 6/3/2017

---

**Result History**  
CULTURE, URINE on 6/6/2017

---

**Reviewed by List**  
[REDACTED]

---

**View SmartLink Info**  
Culture, Urine (Order #308868172) on 6/3/17

# Lab reports: look for susceptibility results

- Antimicrobial susceptibility results
  - Also called “MICs” (“minimum inhibitory concentration”) with “interps” (“interpretation”)
  - Look for interpretations:
    - S = “susceptible”; listed drug can be used to treat
    - I = “intermediate”; listed drug may not be effective treatment
    - R = “resistant”; listed drug can not be used to treat

**Order** CULTURE, URINE [URC] (Order 308868132)

**Ordering Provider**  
Authorizing [REDACTED]

**Acknowledgement Info**

For Placing Order	At 05/17/17 1927	Acknowledged By [REDACTED]	Acknowledged On 05/17/17 1957
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**Order Frequency**

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AUGMENTIN	MIC RESISTANT	Final
CIPROFLOXACIN	MIC RESISTANT	Final
ERTAPENEM	MIC RESISTANT	Final
GENTAMICIN	MIC SUSCEPTIBLE	Final
NITROFURANTOIN	MIC SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC SUSCEPTIBLE	Final

**Lab and Collection**  
CULTURE, URINE on 6/3/2017

**Result History**  
CULTURE, URINE on 6/6/2017

**Reviewed by List**  
[REDACTED]

**View SmartLink Info**  
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# Laboratory evidence for Carbapenemase Production or Resistance Mechanism

- Phenotypic methods for carbapenemase production:
  - Carba NP
  - Metallo- $\beta$ -lactamase testing (e.g., E-test)
  - Modified Carbapenem Inactivation Method (mCIM)
  - Carbapenem Inactivation Method (CIM)
  - Modified Hodge Test (MHT) positive



# Laboratory evidence for Carbapenemase Production or Resistance Mechanism

- Phenotypic methods for carbapenemase production:
  - Carba NP
  - Metallo- $\beta$ -lactamase testing (e.g., E-test)
  - Modified Carbapenem Inactivation Method (mCIM)
  - Carbapenem Inactivation Method (CIM)
  - Modified Hodge Test (MHT) positive
- Molecular methods for resistance mechanism:
  - PCR (for KPC, NDM, OXA-48, IMP, or VIM)
  - Xpert Carba-R (for KPC, NDM, OXA-48, VIM, IMP)

# Example molecular method result

<u>Test Name</u>	<u>Results</u>	<u>Date Reported</u>
Cepheid GeneXpert Carba-R assay		09/06/2018
KPC	KPC gene DETECTED by real time rtPCR.	
IMP	IMP gene NOT DETECTED by real time rtPCR.	09/06/2018
NDM	NDM gene NOT DETECTED by real time rtPCR.	09/06/2018
OXA	OXA-48 gene NOT DETECTED by real time rtPCR.	09/06/2018
VIM	VIM gene NOT DETECTED by real time rtPCR.	09/06/2018

# Example molecular method result

<u>Test Name</u>	<u>Results</u>	<u>Date Reported</u>
Cepheid GeneXpert Carba-R assay		09/06/2018
KPC	KPC gene DETECTED by real time rtPCR.	09/06/2018
IMP	IMP gene NOT DETECTED by real time rtPCR.	09/06/2018
NDM	NDM gene NOT DETECTED by real time rtPCR.	09/06/2018
OXA	OXA-48 gene NOT DETECTED by real time rtPCR.	09/06/2018
VIM	VIM gene NOT DETECTED by real time rtPCR.	09/06/2018



# Exercise 1

What do you think?

<b>Order</b>		CULTURE, URINE [URC] (Order 308868132)	
<b>Ordering Provider</b>			
Authorizing [REDACTED]			
<b>Acknowledgement Info</b>			
For	At	Acknowledged By	Acknowledged On
Placing Order	05/17/17 1927	[REDACTED]	05/17/17 1957
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Priority:	Start:	Process Instructions:	
STAT	05/17/17 1927	** Minimum Specimen Requirements: 25 ML Urine ** **Submit urine in a container with NO preservative** **Use Orange Screw-capped urine cup, White Screw-capped urine tube or Red top tubes**	
<b>Order Frequency</b>			
Antibiotic		Organism	Organism
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AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/AZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final
<b>Lab and Collection</b>			
CULTURE, URINE on 6/3/2017			
<b>Result History</b>			
CULTURE, URINE on 6/6/2017			
<b>Reviewed by List</b>			
[REDACTED]			
<b>View SmartLink Info</b>			
Culture, Urine (Order #308868172) on 6/3/17			
Ordering Provider NPI ID:			

# Exercise 1

What do you think?

Antibiotic		Organism	
		>100,000 cfu/ml enterobacter cloacae	
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final

# Exercise 1

## What do you think?

- Organism
- Susceptibility

HINT: Carbapenem antibiotics include:

- Doripenem
- Ertapenem
- Imipenem
- Meropenem

Antibiotic		Organism	
		>100,000 cfu/ml enterobacter cloacae	
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final

# Exercise 1

What do you think?

- Organism
- Susceptibility

Antibiotic		Organism	
		<b>&gt;100,000 cfu/ml enterobacter cloacae</b>	
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final

# Exercise 1

What do you think?

- Organism

*Enterobacter cloacae*

- Susceptibility

Resistant to ertapenem

Antibiotic		Organism	
		>100,000 cfu/ml enterobacter cloacae	
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final

# Exercise 1

What do you think?

- Organism

*Enterobacter cloacae*

- Susceptibility

Resistant to ertapenem



CRE alert

Antibiotic		Organism	
		>100,000 cfu/ml enterobacter cloacae	
AMP/SULBACTAM	MIC	RESISTANT	Final
AMPICILLIN	MIC	RESISTANT	Final
AUGMENTIN	MIC	RESISTANT	Final
CIPROFLOXACIN	MIC	RESISTANT	Final
ERTAPENEM	MIC	RESISTANT	Final
GENTAMICIN	MIC	SUSCEPTIBLE	Final
NITROFURANTOIN	MIC	SUSCEPTIBLE	Final
PIPERACILLIN/TAZOBACTAM	MIC	RESISTANT	Final
TRIMETH-SULFAMETHOXAZOLE	MIC	SUSCEPTIBLE	Final

# Exercise 2

What do you think?

**Result**

Urine Culture, Routine  
Result 1

Klebsiella pneumoniae Abnormal

25,000-50,000 colony forming units per mL

Antimicrobial Susceptibility

\*\* S = Susceptible; I = Intermediate; R = Resistant \*\*  
P = Positive; N = Negative

MICS are expressed in micrograms per mL

Antibiotic	RSLT#1	RSLT#2	RSLT#3	RSLT#4
Amoxicillin/Clavulanic Acid	R			
Ampicillin	R			
Cefazolin	R			
Cefepime	R			
- Ceftriaxone	R			
Cefuroxime	R			
Cephalothin	R			
Ciprofloxacin	R			
Ertapenem	S			
Gentamicin	S			
Imipenem	S			
Levofloxacin	R			
Nitrofurantoin	I			
Piperacillin	R			
Tetracycline	R			
Tobramycin	R			
Trimethoprim/Sulfa	R			

## Exercise 2

### What do you think?

- Organism
- Susceptibility

#### Result

Urine Culture, Routine

Result 1

Klebsiella pneumoniae Abnormal

25,000-50,000 colony forming units per mL

Antimicrobial Susceptibility

\*\* S = Susceptible; I = Intermediate; R = Resistant \*\*

F = Positive; N = Negative

MICS are expressed in micrograms per mL

Antibiotic	RSLT#1	RSLT#2	RSLT#3	RSLT#4
Amoxicillin/Clavulanic Acid	R			
Ampicillin	R			
Cefazolin	R			
Cefepime	R			
→ Ceftriaxone	R			
Cefuroxime	R			
Cephalothin	R			
Ciprofloxacin	R			
Ertapenem	S			
Gentamicin	S			
Imipenem	S			
Levofloxacin	R			
Nitrofurantoin	I			
Piperacillin	R			
Tetracycline	R			
Tobramycin	R			
Trimethoprim/Sulfa	R			



## Exercise 2

### What do you think?

- Organism  
*K. pneumoniae*
- Susceptibility  
S to Ertapenem  
S to Imipenem

#### Result

Urine Culture, Routine

Result 1

**Klebsiella pneumoniae** Abnormal

25,000-50,000 colony forming units per mL

Antimicrobial Susceptibility

\*\* S = Susceptible; I = Intermediate; R = Resistant \*\*

F = Positive; N = Negative


MICS are expressed in micrograms per mL

Antibiotic	RSLT#1	RSLT#2	RSLT#3	RSLT#4
Amoxicillin/Clavulanic Acid	R			
Ampicillin	R			
Cefazolin	R			
Cefepime	R			
→ Ceftriaxone	R			
Cefuroxime	R			
Cephalothin	R			
Ciprofloxacin	R			
<b>Ertapenem</b>	<b>S</b>			
Gentamicin	S			
<b>Imipenem</b>	<b>S</b>			
Levofloxacin	R			
Nitrofurantoin	I			
Piperacillin	R			
Tetracycline	R			
Tobramycin	R			
Trimethoprim/Sulfa	R			

## Exercise 2

### What do you think?

- Organism  
*K. pneumoniae*
- Susceptibility  
S to Ertapenem  
S to Imipenem

  
**NOT CRE**

### Result

Urine Culture, Routine

Result 1

**Klebsiella pneumoniae** Abnormal

25,000-50,000 colony forming units per mL

Antimicrobial Susceptibility

\*\* S = Susceptible; I = Intermediate; R = Resistant \*\*

F = Positive; N = Negative

MICS are expressed in micrograms per mL

Antibiotic	RSLT#1	RSLT#2	RSLT#3	RSLT#4
Amoxicillin/Clavulanic Acid	R			
Ampicillin	R			
Cefazolin	R			
Cefepime	R			
→ Ceftriaxone	R			
Cefuroxime	R			
Cephalothin	R			
Ciprofloxacin	R			
<b>Ertapenem</b>	<b>S</b>			
Gentamicin	S			
<b>Imipenem</b>	<b>S</b>			
Levofloxacin	R			
Nitrofurantoin	I			
Piperacillin	R			
Tetracycline	R			
Tobramycin	R			
Trimethoprim/Sulfa	R			

## **Criteria to distinguish new from existing cases:**

- **Different organisms/species/carbapenemases are counted as separate events from other organisms/species/carbapenemases.**
- **There is at least a 12-month interval from previous notification event for clinical cases.**
- **A person with a clinical case should not be counted as a screening/surveillance case thereafter (e.g., patient with known infection who later has colonization of GI tract is not counted as more than one case).**
- **A person with a screening case can be later categorized as a clinical case (e.g., patient with positive peri-rectal screening swab who later develops blood stream infection would be counted in both categories).**

# When should isolates be sent to the State Laboratory of Public Health?

- *Enterobacter* spp., *E. coli* or *Klebsiella* spp. resistant to any carbapenem in the absence of carbapenemase resistance mechanism testing
- Identification of CRE producing a carbapenemase other than KPC may be requested for additional testing.

# How to report

- **ELR, Fax or phone to local health department**
- **Local health department will capture case information in Survey Monkey case report form**
  - **Until NCEDSS module is live**

# What will the local health department ask about the case?

- Other healthcare facility exposures (e.g., long-term care facility)?
- International travel/international healthcare?
- Medical devices in place within 2 days prior to culture?
- Wound care?



# What NC DPH is doing:

- **Detect MDROs**

- Increased awareness
- Sentinel surveillance
- Testing at SLPH
- Colonization screening

- **Ensure rapid response & containment**

- Systematic response to even single cases
- Infection prevention assessments
- Inter-facility communication
- Screening for colonization

- **Stewardship efforts**

- Antimicrobial resistance subcommittee
- Get Smart to Be Antibiotics Aware
- STAR partners

- **Education**

- Webinars
- Toolkits
- Presentations
- Guidance documents



# Investigation, containment and response

Goal: contain or slow spread of multidrug-resistant organisms







# **CRE case investigation**

- **Characterize the organism**
  - **Identify if transmission is occurring**
  - **Identify affected patients**
  - **Ensure appropriate control measures are promptly implemented**
-

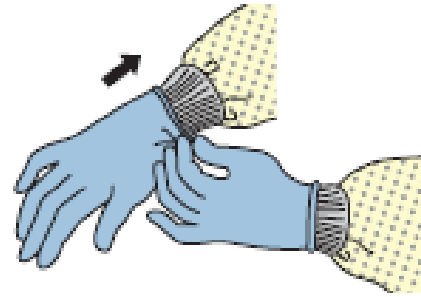
# Prioritize responses given:

1. Organism and mechanism
2. Setting
3. Available resources

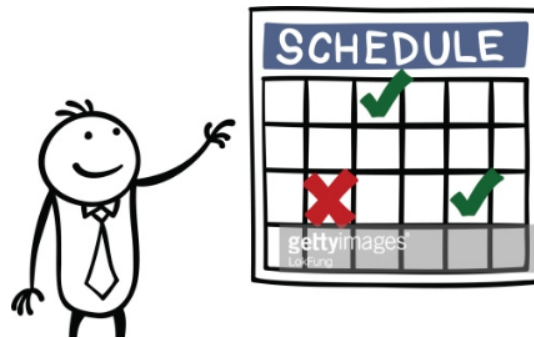
# Standardized response

- **Confirm that a case meets the case definition**
- **Notify patient and healthcare facilities as appropriate**
- **Ensure implementation of control measures**
- **Review the patient's risk factor information**
- **Conduct a healthcare investigation**
- **Contact investigation**
- **Maintain heightened awareness (prospective surveillance) for additional cases in healthcare facility**

# Control measures



Gown and gloves



Prevent opportunities for transmission



Hand hygiene



# Control measures

Environmental cleaning






# Control measures

Communicate CRE status to transferring and receiving facilities

<https://epi.publichealth.nc.gov/cd/hai/docs/InterfacilityTransferInstructionsandForm.pdf>

Transferring Facility Name\*: \_\_\_\_\_  
 Transferring Facility Address\*: \_\_\_\_\_ INTERFACILITY TRANSFER FORM  
 Transferring Facility Phone\*: \_\_\_\_\_ Fax: \_\_\_\_\_  
 Transferred to\*: \_\_\_\_\_ Reason for transfer\*: \_\_\_\_\_  
 Transfer date/time\*: \_\_\_\_\_ / \_\_\_\_\_ / \_\_\_\_\_ Attending physician\*: \_\_\_\_\_ Phone\*: \_\_\_\_\_

Patient/resident demographics and vital signs (date/time taken \_\_\_\_\_ / \_\_\_\_\_)  
 Last Name\*: \_\_\_\_\_ First Name\*: \_\_\_\_\_ DOB\*: \_\_\_\_\_ MRN: \_\_\_\_\_  
 BP\*: \_\_\_\_\_ P\*: \_\_\_\_\_ R\*: \_\_\_\_\_ T(F)\*: \_\_\_\_\_ O<sub>2</sub> SAT\*: \_\_\_\_\_ HT(in): \_\_\_\_\_ WT(lb): \_\_\_\_\_ Diabetic? \_\_\_\_\_ Glucose: \_\_\_\_\_  
 Language  English  Other: \_\_\_\_\_ Mental status\*  Alert  Oriented  Other: \_\_\_\_\_  
 Allergies\*  None  Yes: \_\_\_\_\_ Pain Level (0-10): \_\_\_\_\_ Site: \_\_\_\_\_  
 At risk alerts\*  None  Falls  Aspiration  Pressure ulcers  Seizures  Elopement  Other: \_\_\_\_\_  
 Advanced directives\*  DNR  DNI  MOST  Living Will  Proxy, Contact \_\_\_\_\_

Current isolation precautions\*/required PPE (Check, if indicated)  
 No  Yes, specify  Contact  Droplet  Airborne  
 PPE, specify      

Organisms / infections\*  None  Yes, specify type/date

Multi-drug resistant organisms (MDROs)	Current infection	Hx/Colonized	Pending result
	Date	Date	Date
Methicillin-resistant Staphylococcus aureus (MRSA)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Vancomycin-resistant Enterococci (VRE)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Acinetobacter not susceptible to carbapenems	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Enterobacteriaceae resistant to carbapenems (i.e. CRE)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Extended-spectrum beta-lactamase producer (ESBL)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Clostridium difficile (C. diff)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>
Other: (e.g. Group A Streptococcus (GAS), lice, scabies, disseminated shingles, norovirus, flu, TB, etc.)	<input type="checkbox"/>	<input type="checkbox"/>	<input type="checkbox"/>

Current or recent (last 7 days) symptoms  None  Yes, specify  
 Draining wounds  Concerning rash (e.g. vesicular)  Cough/uncontrolled respiratory secretions  
 Vomiting  Acute diarrhea or incontinent of stool  Other: \_\_\_\_\_

Sensory status and activities of daily living\*

Vision	Hearing	Speech	Ambulate	Transfer	Toileting	Meals	Hygiene	Dressing
<input type="checkbox"/> Good	<input type="checkbox"/> Good	<input type="checkbox"/> Good	<input type="checkbox"/> Self	<input type="checkbox"/> Self	<input type="checkbox"/> Self	<input type="checkbox"/> Self	<input type="checkbox"/> Self	<input type="checkbox"/> Self
<input type="checkbox"/> Poor	<input type="checkbox"/> Poor	<input type="checkbox"/> Difficult	<input type="checkbox"/> Assist	<input type="checkbox"/> Assist	<input type="checkbox"/> Assist	<input type="checkbox"/> Assist	<input type="checkbox"/> Assist	<input type="checkbox"/> Assist
<input type="checkbox"/> Blind	<input type="checkbox"/> Deaf	<input type="checkbox"/> Aphasia	<input type="checkbox"/> Not able	<input type="checkbox"/> Not able	<input type="checkbox"/> Incontinent	<input type="checkbox"/> Tube	<input type="checkbox"/> Not able	<input type="checkbox"/> Not able
Sfy: _____	Sfy: _____				Sfy: _____	Date: _____		

Current devices / recent (last 90 days) procedures\*  None  Yes, specify  
 Tracheostomy tube  Hemodialysis catheter  Procedure, specify type \_\_\_\_\_ and date \_\_\_\_\_  
 Gastrostomy tube  Urinary catheter (date inserted) \_\_\_\_\_  Central line/PICC (date inserted) \_\_\_\_\_

Current medications\*  None  Yes, refer to attached MAR

Vaccination / test history\*  None  Yes, specify

Vaccine/test	Influenza (seasonal)	Pneumococcal	Zoster	Td	Tdap	Tuberculin skin test
Date administered						
Self-report vaccine/test receipt?	<input type="checkbox"/> Yes <input type="checkbox"/> No	<input type="checkbox"/> Yes <input type="checkbox"/> No	<input type="checkbox"/> Yes <input type="checkbox"/> No	<input type="checkbox"/> Yes <input type="checkbox"/> No	<input type="checkbox"/> Yes <input type="checkbox"/> No	<input type="checkbox"/> Yes Result: <input type="checkbox"/> Pos <input type="checkbox"/> No <input type="checkbox"/> Neg

Personal items sent with patient/resident  None  Specify (e.g. glasses, etc.): \_\_\_\_\_

Contact information \_\_\_\_\_

Notes: \_\_\_\_\_

# Containment response elements

	Tier 1 Novel resistance mechanisms, PanR	Tier 2 Mechanisms and organisms not regularly found in a region	Tier 3 Mechanisms and organisms regularly found in a region but not endemic
Infection control assessment	Yes (Green)	Yes (Green)	Yes (Green)
Prospective surveillance	Yes (Green)	Yes (Green)	Yes (Green)
Lab Lookback	Yes (Green)	Yes (Green)	Yes (Green)
Screening of healthcare roommates	Yes (Green)	Yes (Green)	Yes (Green)
Broader screening of healthcare contacts	Yes (Green)	Sometimes (Yellow)	No (Red)
Household contact screening	Yes (Green)	Sometimes (Yellow)	No (Red)
Environmental sampling	Sometimes (Yellow)	No (Red)	No (Red)
Healthcare personnel screening	Sometimes (Yellow)	No (Red)	No (Red)

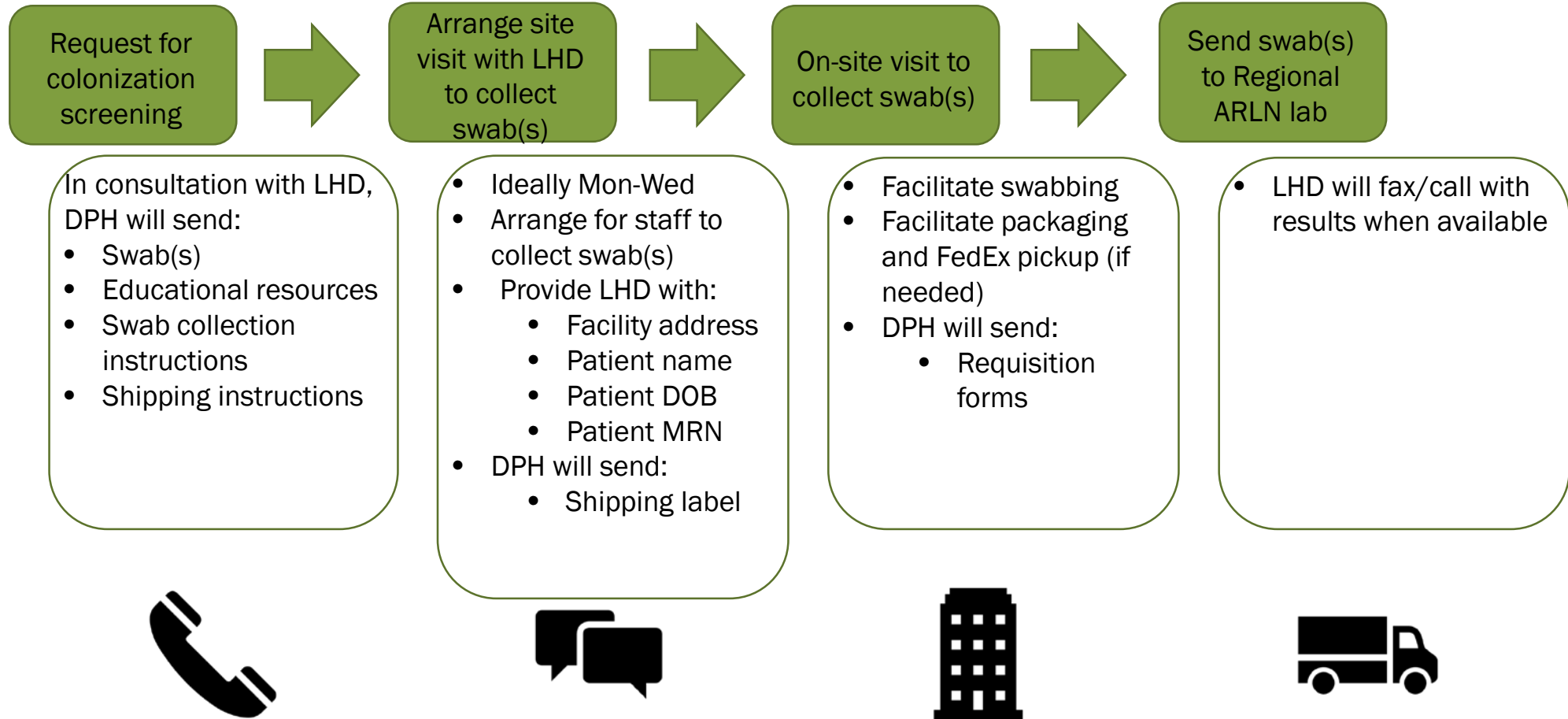
Yes  No  Sometimes 

# Contact investigation

- In consultation with DPH,
  - Screen roommates (and potentially others) that are epidemiologically linked because of healthcare exposure



# Colonization screening process



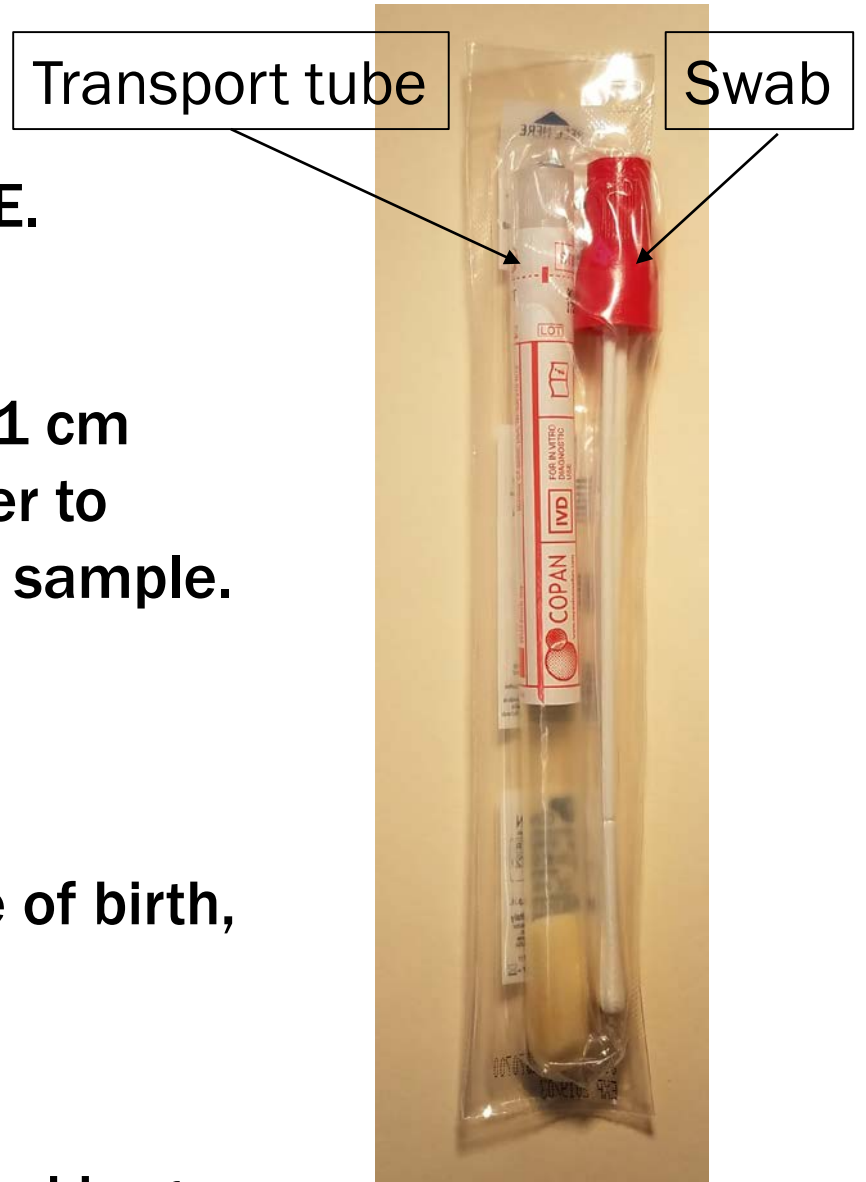
# Swab collection kit

1. Biohazard bag (with absorbent material)
2. Collection instructions
3. Parafilm
4. Swab



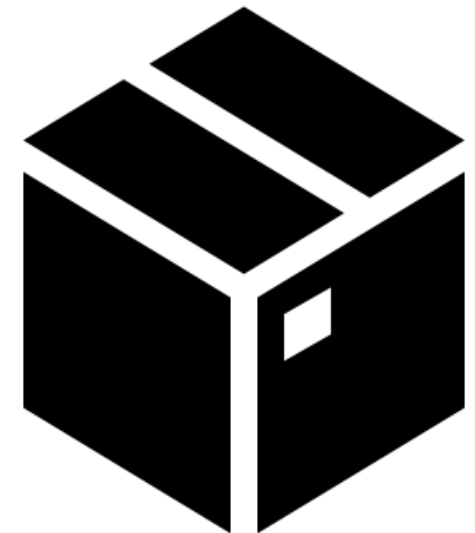
# Swab collection process

1. Perform hand hygiene and apply appropriate PPE.
2. Remove swab from packaging.
3. Carefully insert both tips of swab approximately 1 cm beyond the anal sphincter and rotate gently. Refer to collection instructions for appropriate amount of sample.
4. Uncap transport tube and insert swab.
5. Cap transport tube and seal with Parafilm.
6. Label the transport tube with patient name, date of birth, and collection date.
7. Place transport tube in biohazard bag and seal.
8. Place requisition form in outer pocket of biohazard bag.



# Packing and shipping

Guidance for packing and shipping Category B biological substances via FedEx will be provided prior to swab collection



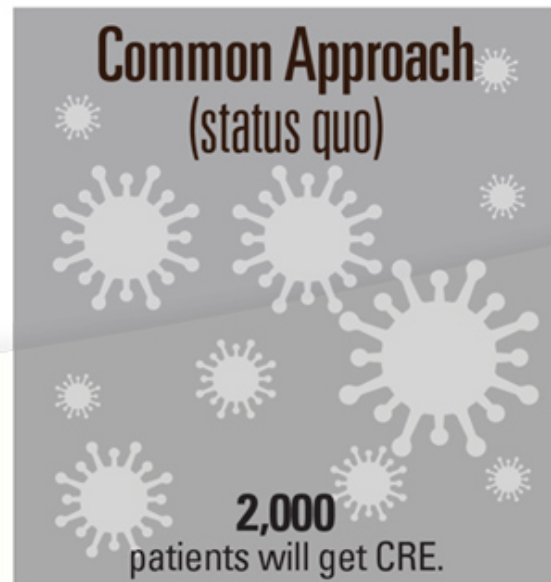
# Partnership is essential

- **CDC & Antibiotic Resistant Laboratory Network (ARLN)**
- **State Laboratory of Public Health (SLPH)**
- **North Carolina Division of Health Service Regulation (DHSR)**
- **Statewide Program for Infection Prevention and Epidemiology (SPICE)**
- **Local Health Departments**
- **Facilities**

# Coordinated approaches prevent MDROs

**More patients get infections when facilities do not work together.**

(Example: 5 years after CRE enters 10 facilities in an area sharing patients)



CRE will impact **12%** of patients.



CRE will impact **8%** of patients.



CRE will impact **2%** of patients.



# How Can My Facility Prevent transmission of MDROs?

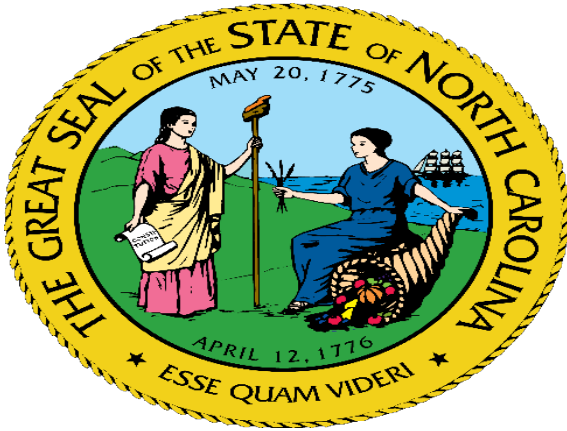
1. Staff education
2. Laboratory notification
3. Cohort residents and staff
4. Contact precautions
5. Hand Hygiene
6. Environmental cleaning
7. Communicate MDRO status
8. Review infection prevention policies and procedures
9. Antimicrobial Stewardship

# How Can My Facility Prevent transmission of MDROs?

- **Communicate with your laboratory**
  - methods for CRE identification
  - capacity to test for CP-CRE
- **Perform periodic reviews of laboratory data**
  - quantify incidence
  - detect changes in overall trends
- **Consider performing rectal screening to detect CRE colonization when admitting patients who have been hospitalized outside the U.S. within the past 6 months**

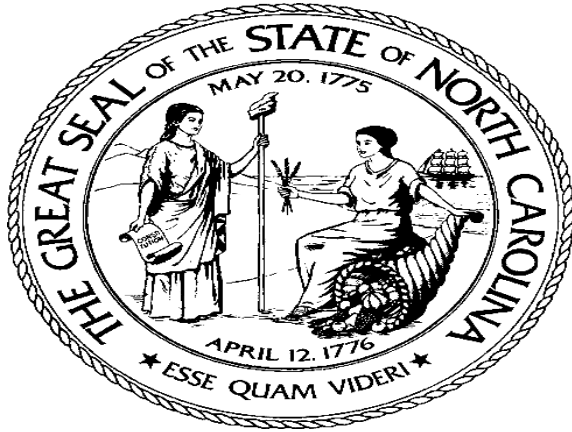


# Questions?



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Epi-On-Call: 919-733-3419





# **CRE Tabletop for Infection Preventionists**

# **PART ONE: NOTIFICATION**

## PART ONE: NOTIFICATION

March 6, 2018

When reviewing your microbiology labs for routine surveillance at acute care hospital A (ACH A), you identify a 63 year old male patient (Mr. Z) with a *Klebsiella oxytoca* isolate from a wound culture taken on March 2.



The isolate is resistant to ertapenem.

No additional testing was performed.

## Is this reportable to NC DPH?

Yes, the result should be reported but the isolate does not need to be sent to the state lab because we confirmed it was CRE **A**

Yes, the result should be reported and the isolate should be forwarded to the state lab **B**

No, *Klebsiella oxytoca* are not covered under the new CRE reporting rule **C**

No, because the only carbapenem the isolate is resistant to is ertapenem **D**

# PART ONE: NOTIFICATION

March 6, 2018

You report the case of CRE to your local health department and the isolate is forwarded to the state lab for resistance mechanism testing.



# PART ONE: NOTIFICATION

March 9, 2018

The health department calls to let you know the state lab results are in. The isolate tests positive for Verona Integron-encoded metallo- $\beta$ -lactamase (VIM). VIM is a plasmid-mediated Carbapenemase that is rare in the U.S. This is the first VIM Producing CRE for the hospital and for the region.



## What does this lab result mean? Choose all that apply

The isolate is a carbapenemase producing CRE (CP-CRE)

Additional case investigation is warranted

Additional public health action is warranted





# What should your facility do to prevent transmission?



# What are some key questions to ask to begin planning your containment strategy?

# Remember!

- You know that in healthcare settings, *Klebsiella* bacteria are primarily spread through person-to-person contact
- This is a novel CRE mechanism so your goal is containment!

## Consider:

1. Where do you go for additional information to plan your response?
2. What are some key questions to ask to begin planning your containment strategy?

## **PART TWO: CASE HISTORY**

## PART TWO: CASE HISTORY

January 1-31 2018 Hospital X – Athens, Greece

On January 1, while on vacation in Greece, Mr. Z suffered a stroke. He was hospitalized in Athens in an ICU.

During his stay he developed a sacral decubitus ulcer. As part of treatment, Mr. Z received several courses of antibiotics.

On January 31, Mr. Z was discharged and flew home with family.



## PART TWO: CASE HISTORY

January 31, 2018 Hospital A – USA

After landing in the U.S., his family brought him directly to the emergency room at Hospital A and he was admitted.

The decubitus ulcer showed no sign of infection. Mr. Z remained bed-bound. He was continent of stool, but required assistance with activities of daily living.

Mr. Z had no roommates during his stay. He was not on contact precautions.



## PART TWO: CASE HISTORY

February 1-28, 2018 Nursing Home 1 – USA

Late on February 1, Mr. Z was transferred to the Nursing Home A. He was admitted to a double occupancy room with Roommate Y.

Mr. Z was not on contact precautions.

On February 23, Mr. Z's wound began to look worse and he developed a fever. He was empirically started on antibiotics. His fever continued and he developed diarrhea.



## PART TWO: CASE HISTORY

March 1-4, 2018 Hospital A – USA

On March 1, Mr. Z was readmitted to the hospital and on March 2, Mr. Z's physician ordered for the decubitus ulcer to be cultured.

On March 4, the CRE result was reported from the clinical laboratory and Mr. Z was placed on contact precautions.





## What risk factors did the patient have for CP-CRE?

Exposure to domestic animals,  
international hospitalizations, recent  
use of antibiotics

International hospitalizations,  
history of smoking, lack of exercise

International hospitalizations, recent  
use of antibiotics, open wound

Open wound, exposure to domestic  
animals, history of smoking


## Why might the LTCF or health department call you about the CP-CRE result?

To blame you for sending a CRE case patient to the LTCF

Notifying facilities where the case had recent healthcare is part of public health response because regional prevention measures are key to control

To brag about being the first facility to identify this mechanism of resistance

Because Mr. Z's friend wrote about it on Facebook



**Consider: Based on the patient's case history, 1. How would you evaluate the potential for transmission? What are key next steps for Hospital A?**

# **PART THREE: FACILITY ASSESSMENT**

## **PART THREE: FACILITY ASSESSMENT**

**After a site visit performed at your hospital in conjunction with the health department, you find multiple issues including:**

- Poor adherence to hand hygiene**
- Patient care equipment stored around inpatient room sinks**
- Improper application/removal of PPE**



## Which of the following are risk factors for CP-CRE transmission? Choose all that apply

Failure to clean physical therapy equipment between patients

Poor staff adherence to hand hygiene

Limited availability of gowns and gloves for contact precautions

Patient care equipment stored around inpatient room sinks

Improper application/removal of PPE

Reusing scissors during wound care

Several infection prevention issues were identified at Hospital A.

How would you:

1. Immediately address the infection prevention concerns?
2. Identify if transmission has occurred?

# **PART FOUR: SCREENING**

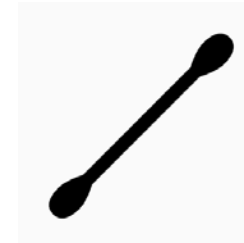


## **PART FOUR: SCREENING**

**March 9, 2018**

**In consultation with the health department,  
your facility screens:**

**5 hospital patients who overlapped with Mr. Z's  
current hospital admission in the same unit for  
three or more days before he was placed on  
contact precautions**



**Mr. Y (Mr. Z's roommate at LTCF 1, who has  
been transferred to your hospital with  
pneumonia)**

# Remember!

**DPH can coordinate colonization screening through our regional antimicrobial resistant laboratory network (ARLN) lab**

**Consider:**

- 1. What data do you want to collect on those you screen?**
- 2. What sites are appropriate for CRE screening?**
- 3. How will swabs be acquired?**

## PART FOUR: SCREENING

March 11, 2018

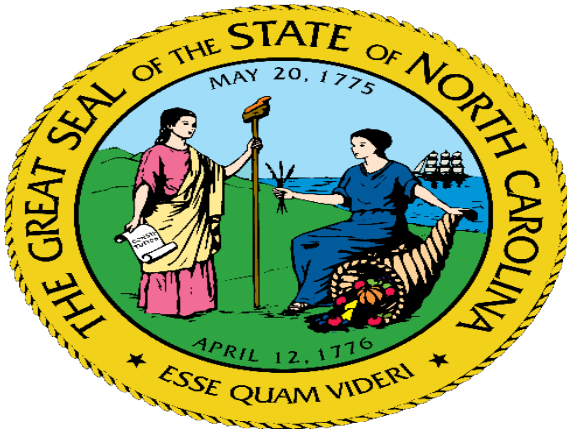
Screening identifies one colonized case of VIM



## **PART FIVE: CONCLUSION**

- **The response continues for 4 months**
- **Screening is expanded to include other high risk patients and continues until there are two consecutive screens with no new positives**
- **Education and rounding continue to ensure appropriate IP practices**
- **Your facility participates in a public health regional training session to educate other hospitals, long-term care facilities, transport services and LTACHS in your area to ensure a coordinated approach to MDRO control**

# Questions?



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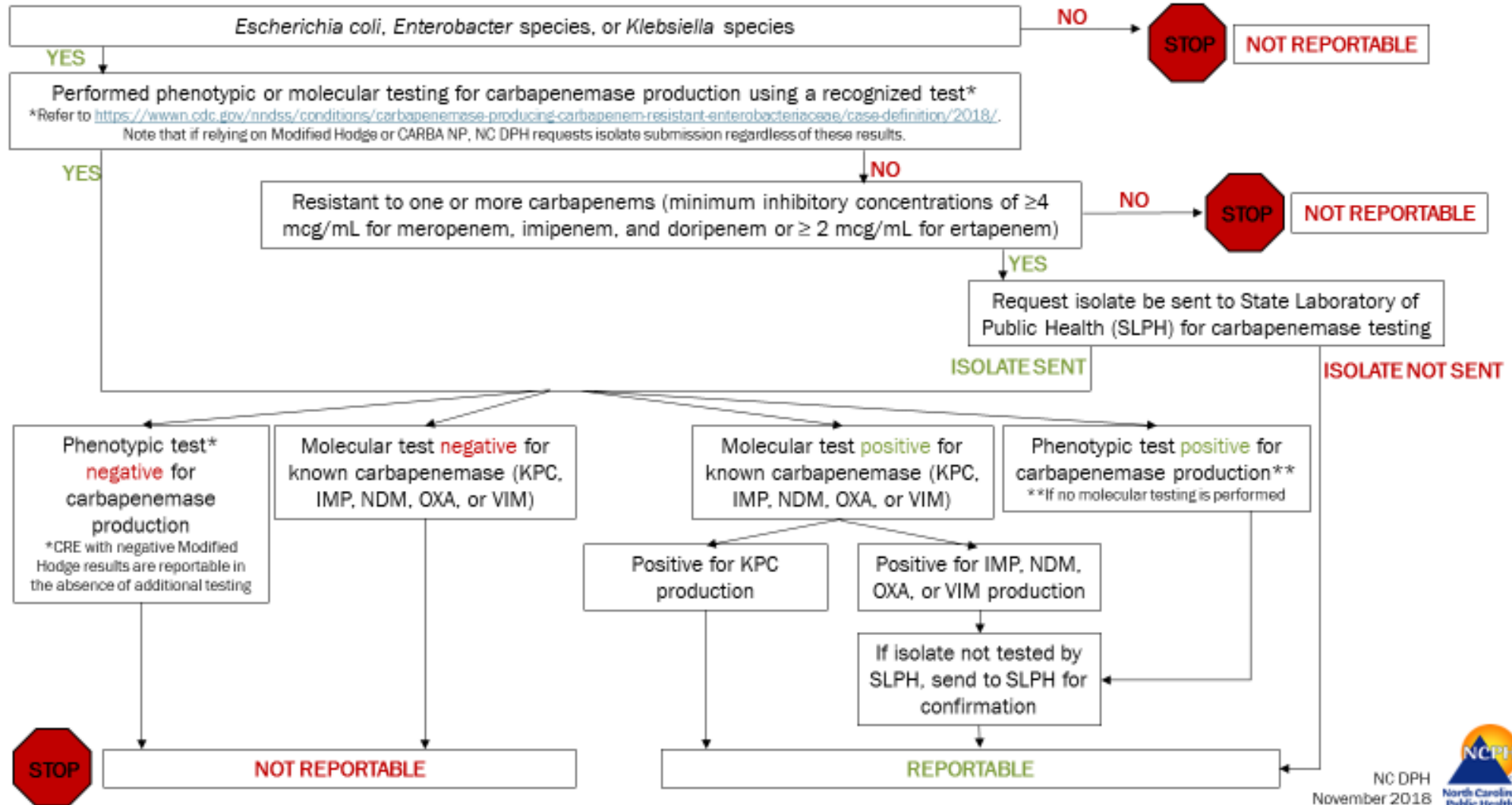


# Resources:

- [https://epi.publichealth.nc.gov/cd/lhds/manuals/cd/reportable\\_diseases.html](https://epi.publichealth.nc.gov/cd/lhds/manuals/cd/reportable_diseases.html)
- **\*\*NEW\*\* Carbapenem Resistant Enterobacteriaceae (CRE)**
  - [Investigation overview](#)
  - [Case Definition](#)
  - [Algorithm for new cases](#)
  - Case report form **\*\*Temporary until NCEDSS is Live\*\***
    - [Carbapenemase-producing carbapenem-resistant Enterobacteriaceae \(CP-CRE\) Case Report Form Survey](#)
  - [CRE Lab Guide](#)
  - Resources:
    - [CDC CRE Toolkit](#)
    - [Interim Guidance for a Public Health Response to Contain Novel or Targeted Multidrug-resistant Organisms \(MDROs\)](#)
    - [MDRO toolkit for long-term care and assisted living facilities](#)
    - [Management of MDROs](#)

# Reporting Algorithm

## Reporting Carbapenemase-Producing Carbapenem-Resistant Enterobacteriaceae (CP-CRE) to NC DPH



# Containment resources:

- **Management of Multidrug Resistant Organisms in Healthcare Settings, 2006**  
[https://www.cdc.gov/hicpac/mdro/mdro\\_to\\_c.html](https://www.cdc.gov/hicpac/mdro/mdro_to_c.html)
- **Interim Guidance for a Public Health Response to Contain Novel or Targeted Multidrug-resistant Organisms (MDROs)**  
<https://www.cdc.gov/hai/outbreaks/docs/Health-Response-Contain-MDRO.pdf>
- **Facility Guide for Control of CRE**  
<https://www.cdc.gov/hai/pdfs/cre/CRE-guidance-508.pdf>
- **Antimicrobial Stewardship**  
<http://epi.publichealth.nc.gov/cd/antibiotics/campaign.html>
- [NCHAI@DHHS.NC.GOV](mailto:NCHAI@DHHS.NC.GOV)





# I understand the process for CRE reporting and surveillance



# I can identify resources and key stakeholders related to CRE response



# I can describe containment strategies for CRE



# I am comfortable assessing infection prevention measures to prevent CRE transmission



# I understand the process for initiating CRE colonization screening

