

#### "MDRO Tiers in Nebraska: Spotlight on Tier 2 Screening" NE APIC Conference, Oct 11, 2024

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## Today's Presentation

- MDRO
- DEFINITIONS
- MDRO TIERS in NEBRASKA
- COLONIZATION SCREENING
- CASE STUDY



Multi-Drug-Resistant-Organisms or MDROs are defined as microorganism, predominantly bacteria, that are resistant to one or more classes of antimicrobial agents

Inappropriate prescribing and use of antibiotics contribute to this growing problem along with other things

https://www.cdc.gov/infection-control/hcp/mdro-

management/background.html

#### Importance of MDROs: Prevention and Control is a National Priority

- MDROs like MRSA, VRE, and certain gram-negative bacteria have significant infection control implications
- MDRO transmission is common in acute care but affects all healthcare settings ranging from ranging from long-term care to specialized units like ICUs and NICUs
- Severity and disease extent vary by population and institution, requiring tailored prevention strategies
- Successful prevention requires administrative and scientific leadership, financial/human resources, and ongoing evaluation

https://www.cdc.gov/infection-control/hcp/mdro-management/introduction.html





#### **Emerging Carbapenem-Resistant Organisms**



Carbapenem-Resistant Enterobacterales Multidrug-Resistant Pseudomonas aeruginosa Carbapenem-Resistant Acinetobacter

https://www.cdc.gov/drugresistance/pdf/threats-report/2019-ar-threats-report-508.pdf



## DEFINITIONS





## • "C" is for CARBAPENEM

• The Carbapenem antibiotics

- Doripenem
- Ertapenem
- Imipenem
- Meropenem



## **Carbapenem Place in Therapy**

- Antibacterial agents with a broad range of antimicrobial activity and a critical place in therapy
- Active against many organisms that are resistant to other β-lactam antibiotics
- Increasingly important due to increase in resistance to other antibiotics
- Relied on to treat sickest patients and most resistant bacteria for over 20 years



The carbapenem antibiotic imipenem



#### Carbapenem Place in Therapy



Utilized for different infections (usually) Pneumonia
Intra-abdominal infection
Urinary tract infections
Meningitis
Skin and soft tissue infections

Off-label use • Most other sites of infection

The Carbapenem antibiotic ertapenem



С-<mark>R</mark>-Е?

### "R" is for RESISTANCE

 Expanded use of Carbapenem has resulted in some Carbapenem resistance in some Gramnegative organisms such as Enterobacterales and Pseudomonas

#### **Mechanisms of Carbapenem Resistance**



Zowawi HM, et al. Nat Rev Urol 2015;12:570-84.



# c-r-<mark>E?</mark>

- "E" is for ENTEROBACTERALES
  - Gram-negative bacteria
  - Found in gastrointestinal tract
  - Cause infection in both healthcare and community settings
- Common Enterobacterales: Klebsiella, Citrobacter, Escherichia coli, Proteus, Enterobacter





What is CP-CRE?

- Carbapenemaseproducing bacteria are more likely to spread their resistance to other bacteria
- "CP" is for CARBAPENEMASE PRODUCING
- CarbapenemASE : Enzymes that break down carbapenems and related antimicrobials making carbapenems ineffective
- The enterobacterales themselves produce this enzyme



#### **CRO versus CPO**

- CRO: Carbapenem-Resistant Organism
  - Any organism resistant to carbapenem antibiotics
  - Not dependent on a carbapenemase
- CPO: Carbapenemase-Producing
   Organism
  - Any organism that produces a carbapenemase
  - A special subset of Carbapenem-Resistant Organisms





#### **CRO versus CPO**

- Carbapenem-Resistant Organisms (CRO)
  - CRAB: Carbapenem-resistant Acinetobacter baumannii
  - CRPA: Carbapenem-resistant Pseudomonas aeruginosa
  - **CRE**: Carbapenem-resistant Enterobacterales
- Carbapenemase Producing Organisms (CPO)
  - CP-CRAB: Carbapenemase-Producing Carbapenem-resistant Acinetobacter baumannii
  - CP-CRPA: Carbapenemase-Producing Carbapenem-resistant Pseudomonas aeruginosa
  - **CP-CRE**: Carbapenemase-Producing Carbapenem-resistant

## The Big FIVE Carbapenemase Genes

**KPC**: *Klebsiella pneumonia* carbapenemase

NDM: New-Delhi Metallo-beta-lactamase

VIM: Verona Integron-Encoded Metallo-beta-lactamase

**IMP**: Active-on-imipenem Metallo-beta-lactamase

**OXA**: Oxacilinase



## **MDRO TIERS**

### **CDC's Containment Guidelines**

Tier 4

**Endemic** MDROs in a region and have been targeted by public health for their clinical significance and potential to rapidly spread

Tier 3

Tier 2

Non-endemic MDROs targeted by facility/region for epidemiologic importance

> MDROs identified in healthcare settings, but **not regularly** identified in the region; and organisms for which **no current treatment** exists

Majority of CPO & *Candida auris* Responses

Tier 1

Novel mechanisms that have never (or very rarely) been identified in the United States and for which experience is extremely limited For accessible version go to https://www.edi.gov/balicontainment/galide/insubtmi

Interim Guidance for a Public Health Response to **Contain** Novel or Targeted Multidrug-resistant Organisms (MDROs)



Updated December 202





Centers for Disease Control and Provention National Center for Disease Description of Center for Ensen Zoongtic Infectious Disea

Source: Interim Guidance for a Public Health Response to Contain Novel or Targeted Multidrug-resistant Organisms (MDROs): Updated December 2022 (cdc.gov)



## Definitions: Epidemiologic Stages

The general pattern of MDRO emergence and spread throughout a geographic area:

- No cases identified
- *Limited spread*: sporadic cases or sporadic clusters of epi linked cases
- *Moderate spread*: cluster or clusters of epi-linked cases
- Advanced spread: Clusters of cases across facilities in different pt. transfer networks
- *Endemicity*: cases are regularly identified in healthcare facilities across the region

## Relationship between Prevention and Response Activities

CDC Interim Guidance for a Public Health Response to Contain Novel or Targeted MDROs



### Multidrug-Resistant Organisms (MDRO) Tiers for Nebraska

Tier	Definition of Included Organisms and Mechanisms	Examples (not all inclusive) of organisms/mechanisms for Nebraska	Transmission-Based Precautions Recommendations
Tier 1	Never (or very rarely) been identified in the United States and for which experience is extremely limited	Novel Carbapenemases	Contact precautions until otherwise recommended by HAI/AR team
Tier 2	Primarily associated with healthcare settings and are not commonly identified in the region (i.e., not been previously identified in the region or have been limited to sporadic cases or small outbreaks), corresponding to "not detected" or "limited to moderate spread" epidemiologic stages. No current treatment options exist (pan not- susceptible) and potential to spread more widely.	Pan-resistant organisms* Candida auris Carbapenemase (e.g., KPC, NDM, OXA-48, VIM, IMP) producing organisms (CPO) • Enterobacterales • Pseudomonas aeruginosa • Acinetobacter Baumannii	Contact Precautions <i>Long-term Care Facilities (LTCF):</i> Enhanced barrier precautions (EBP) recommended for colonized resident(s)**
Tier 3	Include MDROs targeted by the facility or region for epidemiologic importance that have been identified frequently across a region, indicating advanced spread, but are not considered endemic	<ul> <li>Extended spectrum beta-lactamase (ESBL) producing organisms</li> <li>Carbapenem-resistant <i>Enterobacterales</i> (CRE)</li> <li>Carbapenem-resistant <i>Pseudomonas aeruginosa</i> (CRPA)</li> <li>Carbapenem-resistant <i>Acinetobacter Baumannii</i> (CRAB)</li> </ul>	Contact Precautions <i>Long-term Care Facilities (LTCF):</i> Enhanced barrier precautions (EBP) considered for colonized resident(s)**
Tier 4	Endemic in a region and have been targeted by public health for their clinical significance and potential to spread rapidly	<ul> <li>Methicillin-resistant Staphylococcus aureus (MRSA)</li> <li>Vancomycin-resistant Enterococci (VRE)</li> </ul>	Contact precautions per facility risk assessment <i>Long-term Care Facilities (LTCF):</i> Enhanced barrier precautions (EBP) considered for colonized resident(s)**

\* Contact tracing and colonization screening may not be indicated for these organisms

\*\*Contact precautions for acute/active infections or uncontained drainage/secretions

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Updated 5.29.2024 <u>https://dhhs.ne.gov/HAI%20Documents/Nebraska%20MDRO%20Tiers.pdf</u>



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Interim Guidance for a Public Health Response to **Contain** Novel or Targeted Multidrug-resistant Organisms (MDROs) Resource to bookmark!





CDC National Center for Disease Control and Prevention National Center for Emerging and Zoonotic Infectious Diseases

Updated December 2022

**Public Health Strategies** to **Prevent** the Spread of Novel and Targeted Multidrugresistant Organisms (MDROs)

### **Resource to** bookmark!

Accessible Link: https://www.cdc.gov/hai/mdro-guides/prevention-strategy.html



**Control and Prevention** ational Center for Emerging and otic Infectious Disea

## MDRO Tiers for Nebraska (QR Code)

• Resource to bookmark!



## MDRO Cheat Sheet for IPs

<u>Multi-Drug Resistant Organism (MDRO) Cheat</u>
 <u>Sheet for Infection Preventionists</u>



		Multi-Drug Resistant Organism (MDRO) Cheat Sheet for Infection Preventionists				
Type of MDRO	Defintion	Laboratory Evidence to Initiate Transmission-Based Precautions (TBP)	TBP	Duration of TBP	Nebraska Tier <sup>14</sup>	NPHL Specimen Submission 10
Carbapenemase-Producing Carbapenem Resistant Organisms Examples: - OP-CR48: Carbapenemase-Producing Carbapenem- resistant. Acinetaboter baumonsi - OP-CR4: Carbapenemase-Producing Carbapenem- resistant. Parudomons enzginose - OP-CRE: Carbapenemase-Producing Carbapenem-	Any organism that produces a Carbapenemuse	Sentification of a Carlagemenase Gene. Most Cammon Carlagemenase Gene: Most Mark, Oux, RV: Mark, NJP NDM: New Dahl Metallo & Jactanase Other Sites: Caucitanase Other Sites: Caucitanase With Venex Interna Most Dahlagemenase Mit Venex Interna Most Dahlagemenase	Contact Precautions Long-term Care Facilities (LTCF): Enhanced barrier presaubons (EBP) recommended for calonized resident(s) <sup>14</sup> In general, CDC does not recommend screening individuals with a history of CPO colonization or individuals to the history of CPO colonization to inform	Continue isolation indefinitely. In general, screening individuals with a history of colonization or infection with a targeted MDRO with the aim of discontinuing transmission-based precautions is not recommended.	Tier 2	Sabrit II lodate di n-hosas o reference laboratory confirmed carbapenemase-producing Enterobacterials (CPE) or Paradomona eeropinaa (CRPA) or Acinetobacter boumoni (CIAB)
resistant Enterobacterales		IMP: Imipenemase	discontinuation of vertical infection control measures			
Carbagenem Resident Enterchacturales (OR) Organimi: Exhectivita og (E.an) Exhectivita og (E.an) Exhectivita (E.an) Exhectivita (E.an) Exhectivita (E.an) Exhectivita (E.Andree, F. Andref exh Morganetis og (E.Anogani, A. E.) Sarreta og (E. Anogani, A. E.) Sarreta og (E. Anogani, A. E.) Andrea (E. Anogani, A. E.) Anogani, A. E. Anogani, A. E.) Anogani, A. E., Anogani, A. E., Anogani, A. E.) Anogani,	CEL due bacteriu di the Entercholatore deis order hard are rescisito the carbagement antibiotics such as meropenem, ertagenem or intigenem.	Any method of the bacterial family fortendearcasis with succeptibility results that indicate resistance (P) or intermediate (1) to integenem, dorigenem, imperem, and/or morganem. Regarding Sacharia that are intrinsically not succeptible to imperem (a generative spp. Adapted to generative spp.), and succeptible to imperem (a generative spp. Adapted to generative spp.), and succeptible to imperem (a generative spp. Adapted to generative spp.), and succeptible to imperem share than imperem is required.	Cantact Precadon Lang term (or Antilies (LTCF)) Entanced anime precascions (E3) considered for colorised resident(y*	Per failing policy and risk assessment Minimal consideration: duration of hospitalization where this organism was identified.	Tier 3	Enterselectionels: Enterpennen MC 2 3 Jug/ml or imagenen MC 2 3 Jug/ml or imagenen MC 2 3 Jug/ml or on-subspittle by disc diffusion enterban (discus are express beina) On OT simulate the (discus figured) Protess species, Providencia species, and Morganella margani anon susceptible to the simplement susceptible to meropenem and ersperem
Multidrug-Resistant (MDR) Acinetobocter	Gram-negative bacteria that are resistant to several types of antibiotics.	Any Activitation of the standard with the intermediate (i) or hexistate (iii) to at least new drug in at the start here of the following seven categories. 1. Extinded-spectrum cephalosporm (cefepine, certaudime) 3. Aming processing cambraicin, generation (in tobramychi) 4. Carbaparense (integrame, moreopenem, doripanem) 5. Pipercessilly haubotcatem	Contact Precautions Long-term Care Facilities (LTCF): Enhanced barrier precautions (EBP) recommended for colonized resident(s)**	Per facility policy and risk assessment Minimal consideration: duration of hospitalization where this organism was identified.	Tier 3	Acinetobocter boumani: Doripenen 2 4 µg/ml or Imbanem 2 4 µg/ml or Meropenem 2 4 µg/ml or non-susceptible by disc diffusion method and resistant to both celepime and celtasidime at MIC 2 16 µg/ml

## Frequently Used Terms

## **Clinical vs. Colonization**



When organisms cause clinically manifested infections



Acquisition and harboring of organisms by individuals who do not show any signs of infections but are at risk of developing infections.





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## Definition: Colonization Screening

- The use of laboratory testing to *identify colonized individuals* by testing for the presence of novel or targeted MDROs on or in the body of an individual.
- When an emerging MDRO is identified, colonization screening is *recommended* by CDC as an essential component of the public health response.
- Colonization screening identifies unrecognized carriers so that infection prevention and control measures can be targeted to prevent the spread of antimicrobial resistance.

 <u>CDC Interim Guidance for a Public</u> <u>Health Response to Contain Novel or</u> <u>Targeted MDROs</u>



## Definitions: Point Prevalence survey (PPS)

- Colonization screening performed unit- or facility-wide following the identification of a patient or resident with a novel or targeted MDRO
- The goal of these surveys is to identify colonized individuals and initiate transmission-based precautions
  - Serve as both an assessment tool (for possible transmission) and an intervention (facilitating identification of colonized individuals for implementation of appropriate precautions)
- <u>CDC Interim Guidance</u>
   <u>for a Public Health</u>
   <u>Response to Contain</u>
   <u>Novel or Targeted</u>
   <u>MDROs</u>



## CPO and *C. auris* Screening – Key to Strategies for Stopping Spread

	R Contraction of the second se			
	Response (Containment)	Prevention		
R Screening	<ul> <li>Identification of an uncommon novel MDRO in a region</li> <li>Suspected or confirmed transmission of a novel MDRO in a healthcare facility (i.e., an outbreak)</li> </ul>	<ul> <li>Based on facility characteristics and local epidemiology, NOT on case or epi links to known case</li> <li>Recurring basis at influential facilities</li> <li>Ad hoc basis at facilities with unknown epidemiology (typically long-term care)</li> </ul>		
Timing	Not predictable (happens when detections occur)	Predictable (screening should be scheduled far in advance)		
Follow-up screening (if cases detected)	Continues until transmission <b>controlled</b> ( <b>stopped or reduced</b> )	Generally, should adhere to fixed schedule (i.e., should not trigger multiple follow up rounds of screening)		

**Contact Isolations:** A set of infection control practices recommended by the CDC to prevent the spread of infectious agents, particularly those transmitted through direct or indirect contact with an infected individual or contaminated surfaces.



#### Enhanced Barrier Precautions

An infection control designed to reduce transmission of multidrug-resistant organisms (MDROs) in nursing homes<sup>\*</sup>

Involve gown and glove use during high-contact resident care activities for residents known to be colonized or infected with a MDRO as well as those at increased risk of MDRO acquisition (e.g., residents with wounds or indwelling medical devices).

\*Individuals may not have known MDRO but may still be in EBP precautions because of indwelling device, chronic wound etc.





#### **CPO Transmission in Healthcare Facilities**



Transmission: When a pathogen moves from one host to another

#### Whole Genome Sequencing (WGS)

The information encoded in the genomes of disease-causing bacteria, viruses, and fungi represent unique genetic fingerprints.

#### Whole Genome sequencing

(WGS) is a laboratory procedure that determines the order of all, or most, of the nucleotides in the genome of these disease-causing microbes.

A nucleotide polymorphism, or SNP (pronounced "snip"), is a variation at a single position in a DNA sequence among individuals



Spotlight: Colonization Screening

## **Screening of Healthcare Contacts**

In general, the recommendations apply to all inpatient healthcare exposures of the index patient in the <u>30 days</u> <u>prior</u> to the identification of the target organism to the present. Depending on the type of exposure and organism, contact investigations may sometimes include healthcare facilities where the patient received care but did not stay overnight

- outpatient clinics
- community contacts

If the index patient had recent inpatient healthcare exposure, screen epidemiologically linked patients.

Screening should occur even if the index patient was being managed with Contact Precautions or Enhanced Barrier Precautions

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Response Elements	Tier 2
Contact Investigation (Typical review period: 30 days prior to culture collection to present)	
Screening of healthcare contacts (i.e., residents and patients)	<mark>ALWAYS</mark>
Household contact screening	RARELY
Healthcare personnel screening	RARELY
Additional Actions if Transmission Identified in Healthcar	e
Recurring response-driven point prevalence surveys <sup>3</sup>	<mark>ALWAYS</mark>
Evaluate potential spread to healthcare facilities that regularly share patients with the index healthcare facility <sup>4</sup>	USUALLY
Clinical Laboratory Surveillance	
Retrospective lab surveillance <sup>6</sup>	ALWAYS
Prospective lab surveillance⁵	ALWAYS
Environmental Cultures	
Environmental sampling	RARELY
Infection Control Measures	
Notify healthcare providers; promptly implement appropriate transmission-based precautions	<mark>always</mark>
Infection control assessment with observations of practice	<mark>ALWAYS</mark>
Clear communication of patient status with transferring facilities	ALWAYS

Tier-2 Containment Response Elements for Infection Preventionists

## **Screening of Healthcare Contacts**





- Roommates and patients who shared a bathroom with the index patient. Screen these contacts even if they have been discharged from the facility to another inpatient setting.
- If discharged to home, consider notifying the contact and offering screening or *flagging the chart to facilitate* preemptive Contact Precautions and *admission screening if they are readmitted in the next six months.*
- Screen the patient currently admitted to room(s) and bed spaces where the index patient stayed at least one night in healthcare facilities identified during the healthcare investigation
- If these individuals have been discharged to high-acuity post-acute care, health departments should consider screening these individuals



In most situations, Broader screening using point prevalence surveys (PPS) is preferred.



Alternatively, broader screening may initially target contacts who are at higher risk of MDRO aquation :

Overlap on the same ward as the index patient Bedbound, High levels of care, Receipt of antimicrobials, or Mechanical ventilation), and Who are still admitted

*Considerations* – When deciding whether to use a risk-factor-based approach, PPS, or both strategies in <u>combination</u>, consider individual facility characteristics, local epidemiology, characteristics of index patient, feasibility of identifying contacts, and laboratory capacity.

If it will take several days to identify higher risk contacts or if most higher risk contacts have been discharged from a facility, perform a unit-wide point prevalence survey promptly.

Screening of Healthcare Contacts continued



#### Ongoing Transmission: How is it different?

Interim Guidance for a Public Health Response to Contain Novel or Targeted Multidrug-resistant Organisms (MDROs)

- When transmission is identified for Tier 2, periodic (e.g., every two weeks) PPS are recommended until transmission is controlled
- Control of transmission
  - Two consecutive PPS with no new MDRO cases identified
  - Substantially decreased transmission when colonization pressure is high
- If transmission does NOT decrease across multiple point prevalence surveys
  - Consider pausing or increasing the interval between PPS (e.g., 4-6 weeks)
  - Reassess and implement measure to improve infection control
- Implement measures to prevent outbreaks at downstream facilities



# Why is colonization screening important

- If we identified one CP-CRE in our facility, how many more do we have?
- Why do we need to know it?

In the next few slides, we will explore a hypothetical scenario to understand this better

- Benefit of knowing ("are there more CP CREs in our facility?") for facilities and patients,
- Benefit of NOT knowing ("are there more CP CREs in our facility?") for facilities and patients

## Case Study/Scenario



- Ms. Blanch, an 80-year-old nursing home resident, can do all her daily activities on her own, walks with a walker, dines and plays bingo with others in her nursing home and goes to her physical therapy on a regular basis.
- She lived in the nursing home named "Road to Eden"
- She developed UTI symptoms a couple of days ago, visited an outpatient and her urine culture tested positive for *E coli* having a resistant Carbapenemase gene (NDM). She was then admitted to a hospital, not returning to her nursing home.
- The lab notified the state HAI/AR program, the program notified both nursing homes and discussed response components including colonization screening
- Note: Ms. Blanch lived in the Road to Eden in the last 30 days prior to her positive culture



# Scenario continues.....



#### One unknown fact



The same nursing home has a second patient with exactly the same *E col*i-NDM:



Mr. T is colonized (not infected) but his colonization status is unknown to all

GOAL of the COLONIZATION SCREENING is to DISCOVER/IDENTIFY the UNKNOWN COLONIZED CASES



Mr. T was in the Bingo Team with Ms. Blanch



#### Title: MDRO Transmission Investigation at Nursing Home: The Story of Mr. T





MDRO Transmission at Nursing Home: The Story of Mr. T Initial Screening: Limited Action. Screened 1 roommate and bingo partner, both tested (-)



Week 2: Mr. T's positive status is unknown, he and his buddies are practicing Taekwondo



2<sup>nd</sup> month: 2 nd PPS (-), 3<sup>rd</sup> PPS 1 (+), (total 8 + in EBP), 2 more PPS to go and continues until 2 consecutive negatives



Outbreak identified: WGS Confirms Transmission

2 more positives (total 7 are in EBP), 2 more rounds of PPS



1<sup>st</sup> month: E. Coli NDM in Mr. T, hospitalized

Screened 10, 4 tested positive, PPS for all 35 started

3<sup>rd</sup> month: Continued Screening and New Positive Cases Escalation to Hospital: Since Mr. T's MDRO status was unknown, when hospitalized, no contact precautions, exposed other patients, screening at ACH

Outcome: 2 facilities, multiple patients Conclusion: Importance of Timely Screening

## Take Aways





INTIMAL AND TIMELY DECISION ABOUT COLONIZATION SCREENING MADE ALL THE DIFFERENCES LEADERSHIP IS IMPORTANT



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# Other screening: Screening process in Nebraska

- Admission screening
- Discharge screening (optional)
- Baseline (Prevention based PPS)



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## Process of Colonization Screening in Nebraska



Select patients to screen

Members of HAI Team and the IP at the facility will identify appropriate epidemiologic contacts for screening based on above guidance.

Order supplies

MN ARLN (Regional CDC laboratory in MN) NPHL (Nebraska State Public Health Lab in Omaha) Facility will screen their patient following the instruction

**Actual screening** 



Mail back for testing

**Receive results** 

## **Healthcare and Household Screening**

#### Healthcare personnel screening:

- □ In the absence of known or suspected transmission from HCP or other strong epidemiologic links, HCP screening is not recommended.
- □ Household contact screening:
  - Screen household contacts who have extensive contact (e.g., share a bed or assist with personal care) with the index patient if the household contact has frequent inpatient healthcare exposure to determine if transmissionbased precautions are necessary for their subsequent admissions.
  - Consider screening other household contacts if household transmission is suspected.
  - □ If household contacts are HCP, prior to pursuing screening consider what actions will be taken if they are colonized (e.g., work restrictions and rescreening).

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## **QUESTIONS?**







## THANK YOU

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