



Michigan Department of Health & Human Services

Michigan Hepatitis A Outbreak Update for Clinicians

Eden V. Wells, MD, MPH, FACPM
For a Clinical Webinar Event

May 7, 2018

Putting people first, with the goal of helping all Michiganders lead healthier and more productive lives, no matter their stage in life.

Disclosures

Dr. Wells has no financial interest, or any conflicts of interest, regarding the material provided in her presentation today

Slide Material has been Adapted from:

- *Epidemiology and Prevention of Vaccine-Preventable Diseases*, National Center for Immunization and Respiratory Diseases
- Michigan Department of Health and Human Services Communicable Disease Division materials

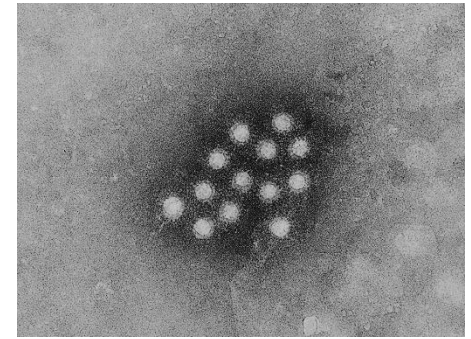
Outline

- Review of Hepatitis A Virus
- Hepatitis A Diagnosis and Management
- Guidelines for Hepatitis A Prevention, General
- Michigan Outbreak 2016-Present
- Guidelines for Management of Hepatitis A, Michigan 2018
- Waning Hep A Immunity in HIV, Michigan 2018

Review of Hepatitis A Virus

Hepatitis A Virus

- Picornavirus (RNA)
- Humans are only natural host
- Stable at low pH
- Inactivated by temperature of 185°F or higher, formalin, chlorine
 - Disinfection: 1 and 2/3 cups bleach in 1 gallon water (5000 ppm).
 - Allow 1 minute of contact time and then rinse with water.



CDC: PHIL 2739

Centers for Disease Control and Prevention. Epidemiology and Prevention of Vaccine-Preventable Diseases. Hamborsky J, Kroger A, Wolfe S, eds. 13th ed. Washington D.C. Public Health Foundation, 2015.

Hepatitis A Pathogenesis

- Entry into mouth
- Viral replication in the liver
- Virus present in blood and feces 10-12 days after infection
- Virus excretion may continue for up to 3 weeks after onset of symptoms

Hepatitis A Disease

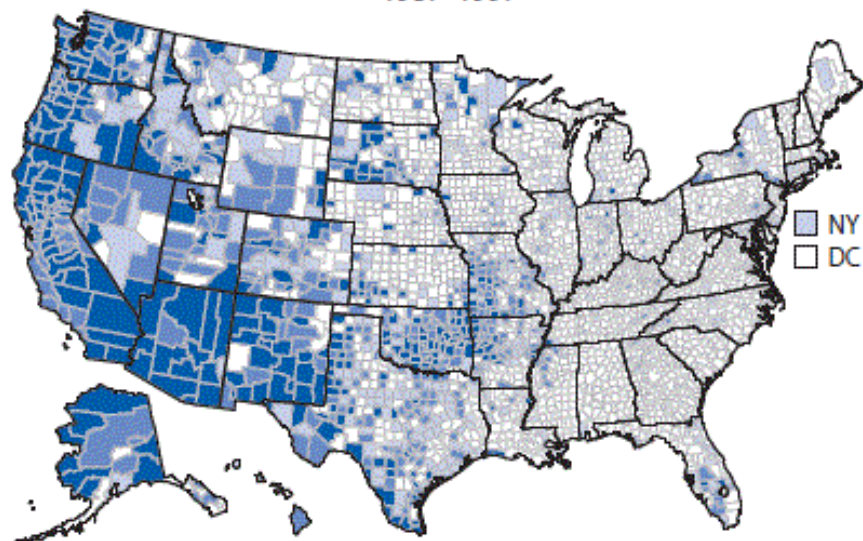
- Highly contagious, vaccine-preventable (acute) liver infection
- Incubation Period - illness can appear **15 to 50** days after exposure
 - **Average 28 days**
- Illness not specific for hepatitis A
- Likelihood of symptomatic illness directly related to age
- Children generally asymptomatic, adults symptomatic

Hepatitis A Epidemiology

- Reservoir
 - human
- Transmission
 - fecal-oral
- Temporal pattern
 - none
- Communicability
 - 2 weeks before illness to 1 week after onset of jaundice

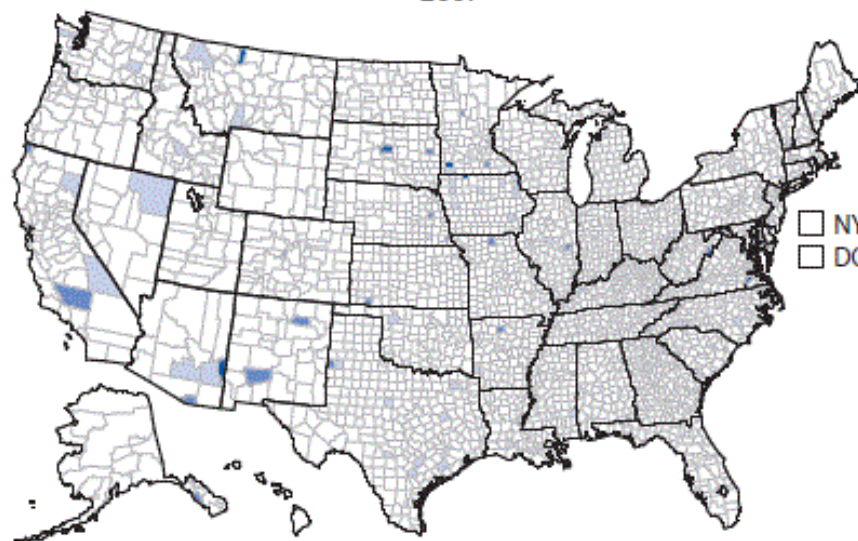
Incidence* of reported acute hepatitis A cases — National Notifiable Diseases Surveillance System, United States,

1987–1997



□ 0-4 □ 5-9 □ 10-19 ■ ≥20

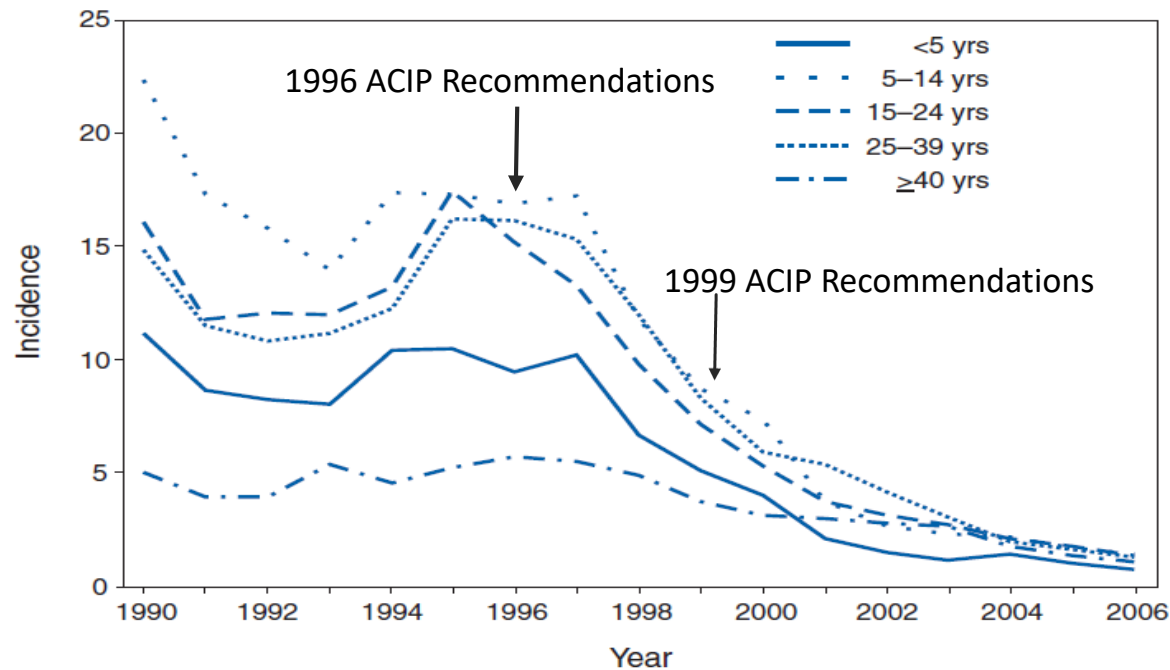
2007



□ 0-4 □ 5-9 □ 10-19 ■ ≥20

* Rates per 100,000 population; † Annual Average Incidence; Source: MMWR Supplements; February 12, 2016 / 65(1);29–41

USA Incidence of acute hepatitis A— by age group: 1990 - 2006



* Per 100,000 population.

Transmission

**Ingestion of fecal matter, *even in microscopic amounts*,
from:**

Touching objects or
eating food that
someone with hepatitis
A infection handled

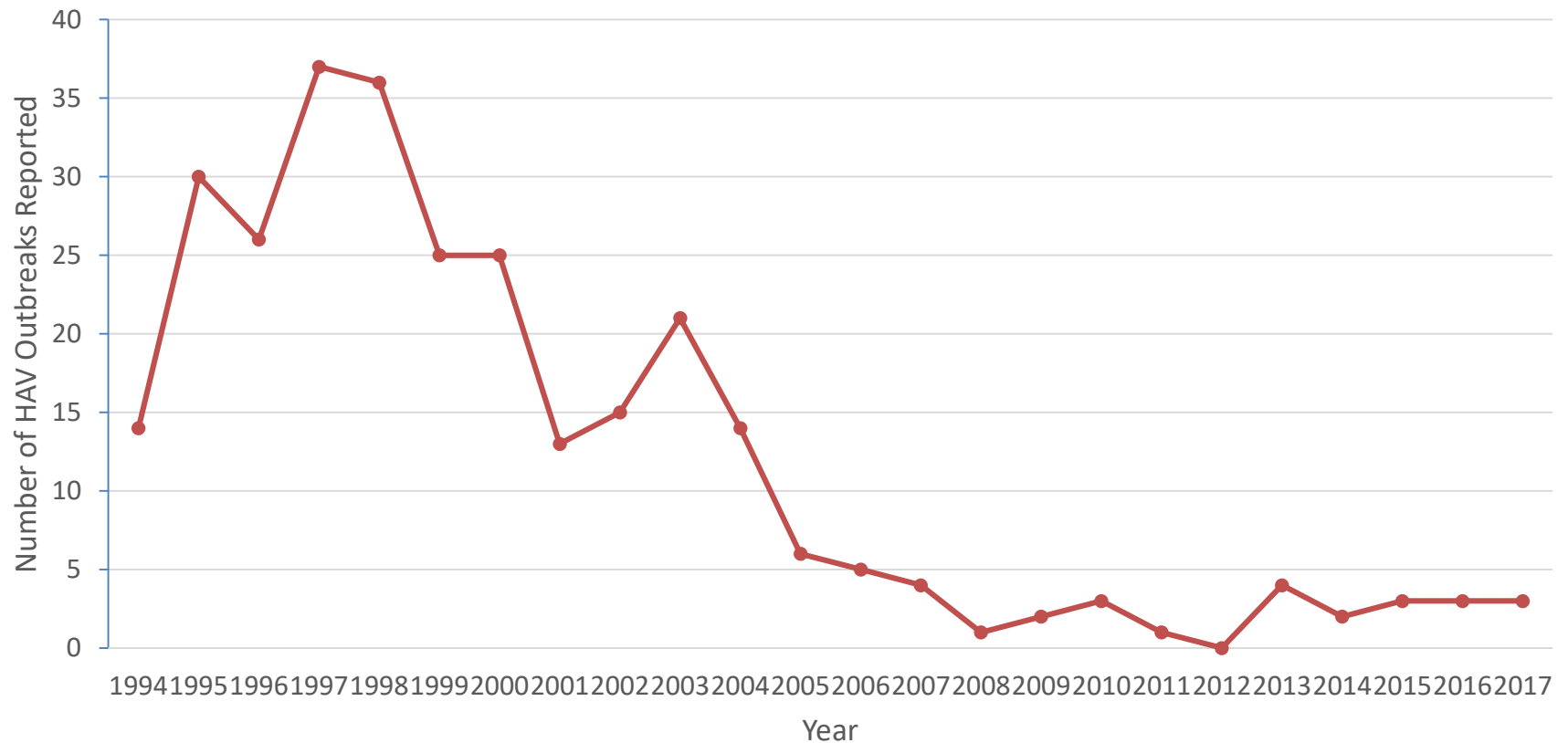
Close, person-to-
person contact with a
person who is infected

Use of recreational
drugs, whether injected
or not

Sexual contact with
someone who has a
hepatitis A infection



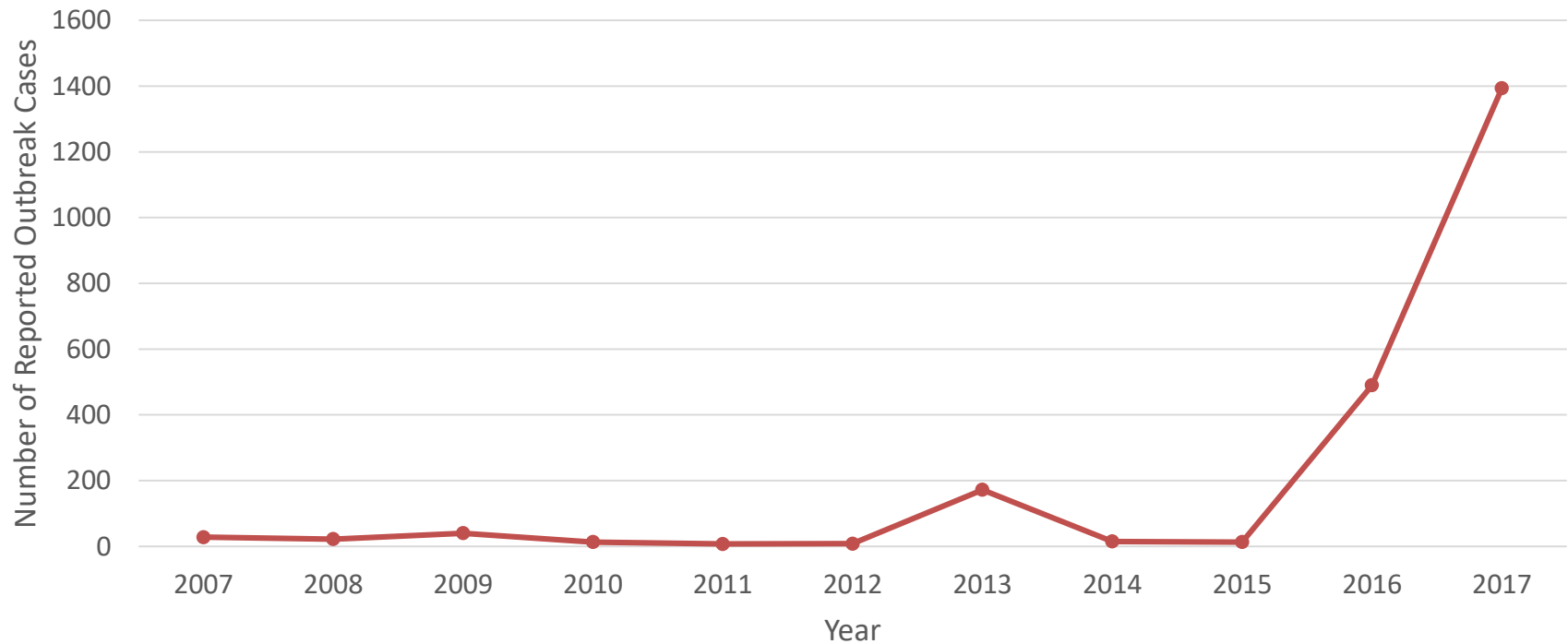
Hepatitis A Outbreaks Identified in the United States – 1994–2017



Craig AS, et al. *Am J Med Sci* 2007 CDC FoodTool

From MDHHS Communicable Disease Hep A Slide set

Reported Cases Associated with HAV Outbreaks – United States, 2007–2017



Craig AS, et al. *Am J Med Sci* 2007; Collier MG, et al. *Lancet Infect Dis* 2014.;
CDC Unpublished data

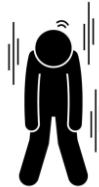
Hepatitis A Diagnosis and Management

Hepatitis A Symptoms

NON-SPECIFIC!!!



Fever



Fatigue



Nausea



Loss of Appetite



Jaundice



Stomach Pain



Vomiting



**Dark Urine, Pale Stools,
or Diarrhea**

Not all people infected with hepatitis A experience illness. Most hepatitis A infections in children younger than age 6 are not accompanied by symptoms. Older children and adults are at risk for severe hepatitis A disease.

Risk Factors for Hepatitis A (General)

- International travelers (particularly high-risk itineraries like travel to rural areas in high-risk countries)
- Contacts of recent international adoptees from HAV endemic countries
- Men who have sex with men
- Users of illegal drugs
- ***Michigan Outbreak 2016-present are different!!***

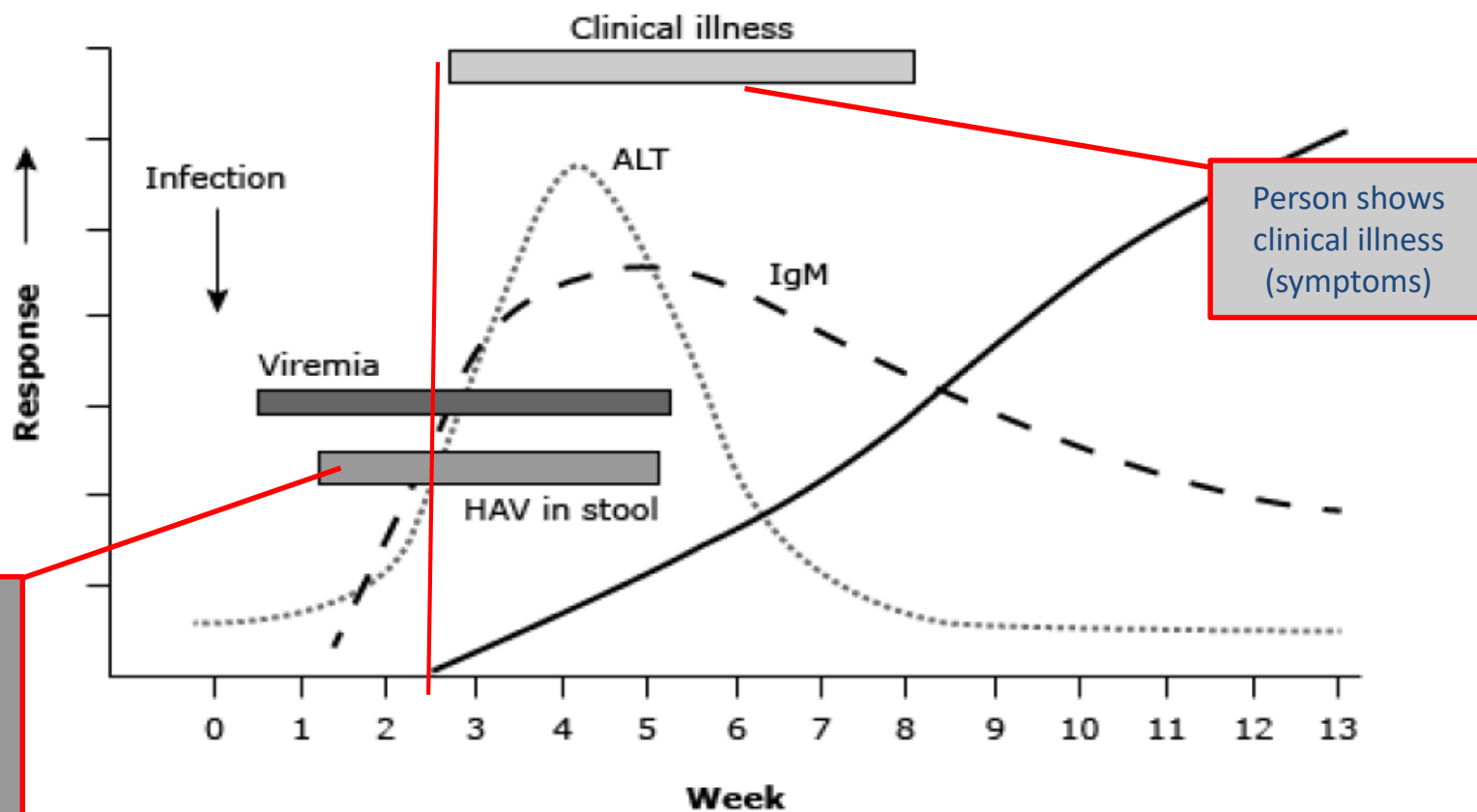
Occupational Risks

- Outbreaks of hepatitis A have been reported among persons working with hepatitis A-infected primates
 - **This is the only occupational group known to be at increased risk for hepatitis A**
- Food workers are not at increased risk because of their occupation but may play a critical role in transmission
- US serologic studies have shown no or mildly increased risk of HAV infection in wastewater workers

Serologic Testing

- Serologic testing required to confirm the diagnosis.
- Virtually **all** patients with acute hepatitis A have detectable IgM anti-HAV.
- Acute HAV infection confirmed during acute or early convalescent phase of infection by presence of serum IgM anti-HAV
 - IgM detectable 5-10 days before the onset of symptoms and can persist for up to 6 months
- Polymerase chain reaction (PCR)-based assays can be used to amplify and sequence viral genomes
 - These assays are helpful to investigate common-source outbreaks of hepatitis A.

Course of hepatitis A



Timeline for hepatitis A manifestations.

Serologic Testing, cont

- IgG anti-HAV appears in the convalescent phase of infection
 - Remains present in serum for the lifetime of the person, and confers enduring protection against disease**
 - Total anti-HAV measures both IgG anti-HAV and IgM anti-HAV
 - Persons with total anti-HAV positive and IgM anti-HAV negative: indicates immunity consistent with either past infection or vaccination

***Exceptions noted- will be discussed in HIV later in presentation*

Medical Management

- There is no specific treatment for hepatitis A virus infection
- Treatment and management of HAV infection are supportive



Guidelines for Hepatitis A Prevention- General

Primary Prevention- Immunization

- Inactivated whole-virus vaccines
- Pediatric and adult formulations
 - pediatric formulations approved for persons 12 months through 18 years
 - adult formulations approved for persons 19 years and older

Hepatitis A Vaccine Immunogenicity

- Adults
 - more than 95% seropositive after one dose
 - nearly 100% seropositive after two doses
- Children and Adolescents
 - more than 97% seropositive after one
 - 100% seropositive after 2 doses (in clinical trials)
- **Hepatitis A Vaccine Efficacy**
 - HAVRIX
 - 40,000 Thai children 1 to 16 years of age
 - vaccine efficacy 94%
 - VAQTA
 - 1,000 New York children 2 to 16 years of age
 - vaccine efficacy 100%

Childhood Hep A Vaccination

- All children should receive hepatitis A vaccine at 12 through 23 months of age
- Vaccination should be integrated into the routine childhood vaccination schedule
- Children who are not vaccinated by 2 years of age can be vaccinated at subsequent visits
- States, counties, and communities with existing hepatitis A vaccination programs for children 2 through 18 years of age should maintain these programs
- New efforts focused on routine vaccination of children 12 months of age should enhance, not replace ongoing vaccination programs for older children
- In areas with without an existing hepatitis A vaccination program catch-up vaccination of unvaccinated children 2 through 18 years of age can be considered

Adult Hep A Vaccination

- Adults 19 years of age and older receive the adult formulation of hepatitis A vaccine according to licensed schedules
- Persons at increased risk for HAV infection, or who are at increased risk for complications of HAV infection, should be routinely vaccinated
 - See “Risk Factors” Slide 16

Post Exposure Prophylaxis (PEP)

- PEP can protect susceptible (unvaccinated) persons who have *recently* been exposed to hepatitis A:
 - Remember, the incubation period of hepatitis A is approximately 28 days (range 15-50 days)
- **PEP must be given within 2 weeks after exposure to prevent infection:**
 - Hepatitis A Vaccine
 - Immune globulin (IG)



Photo credit: National Institute of Allergy and Infectious Diseases (NIAID), 2014. Public Health Image Library.

Michigan Outbreak 2016-Present

Hepatitis A Outbreak in Michigan: Statewide Update

May 4, 2018



Michigan Hep A Outbreak

CURRENT MICHIGAN OUTBREAK

August 2016, nine cases of hepatitis A reported in SE Michigan counties.

Early investigation focused on ill food workers and food establishments

No common source of infected identified; multi-modal

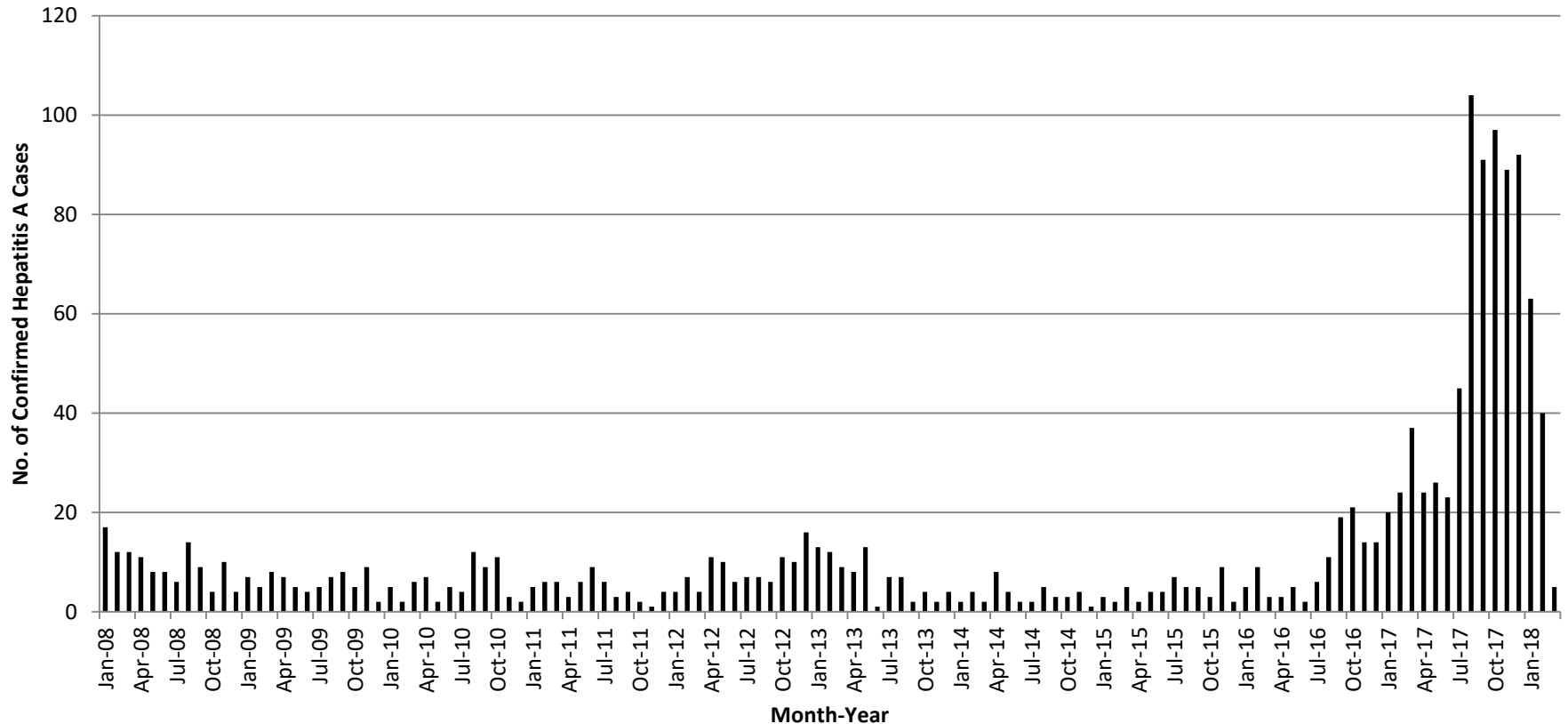
Cases included persons with substance use disorder, homeless or transient living, recently incarcerated, food workers, and men who have sex with men (MSM)

Investigations continuing, vaccination efforts ongoing

Outreach and education to vulnerable populations

8

Reported Number of HAV cases in Michigan 2008-2018



Hepatitis A MI Outbreak Testing

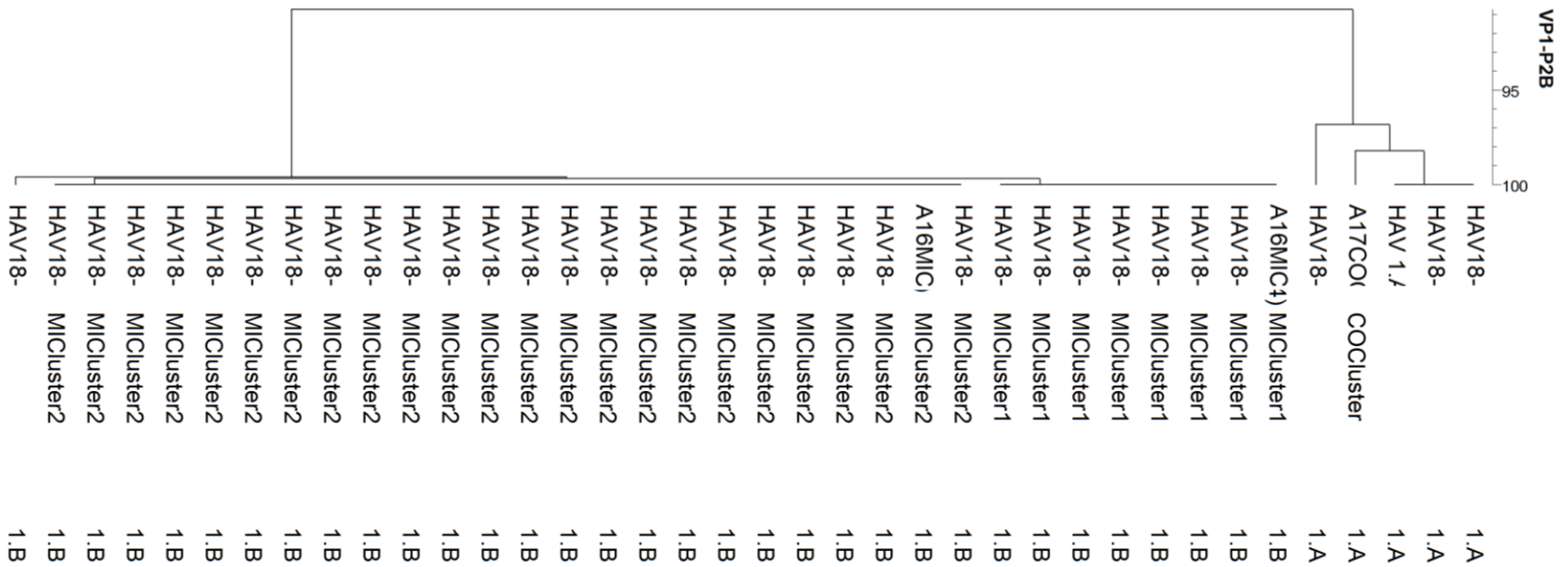
- CDC sequencing HAV positive serum samples from representative cases (high risk cases, but no travel-related cases)
 - Fall 2016, SEMI clinical labs asked to send all HAV positive serum samples to Bureau of Labs (BOL)
 - Samples forwarded to CDC lab for sequencing
 - CDC notified MDHHS in Aug 2016 that MI cases not related to previous hepatitis A outbreaks in Virginia or Hawaii, San Diego
 - Michigan strains **are unique**: 1B strain 1 and 2

Outbreak Case Classification:

- Cases were classified as outbreak cases if they had the Hepatitis A1B Outbreak Strain #1 or #2

BOL began conducting whole genome sequencing Dec 2017

WHOLE GENOME SEQUENCING DENDROGRAM



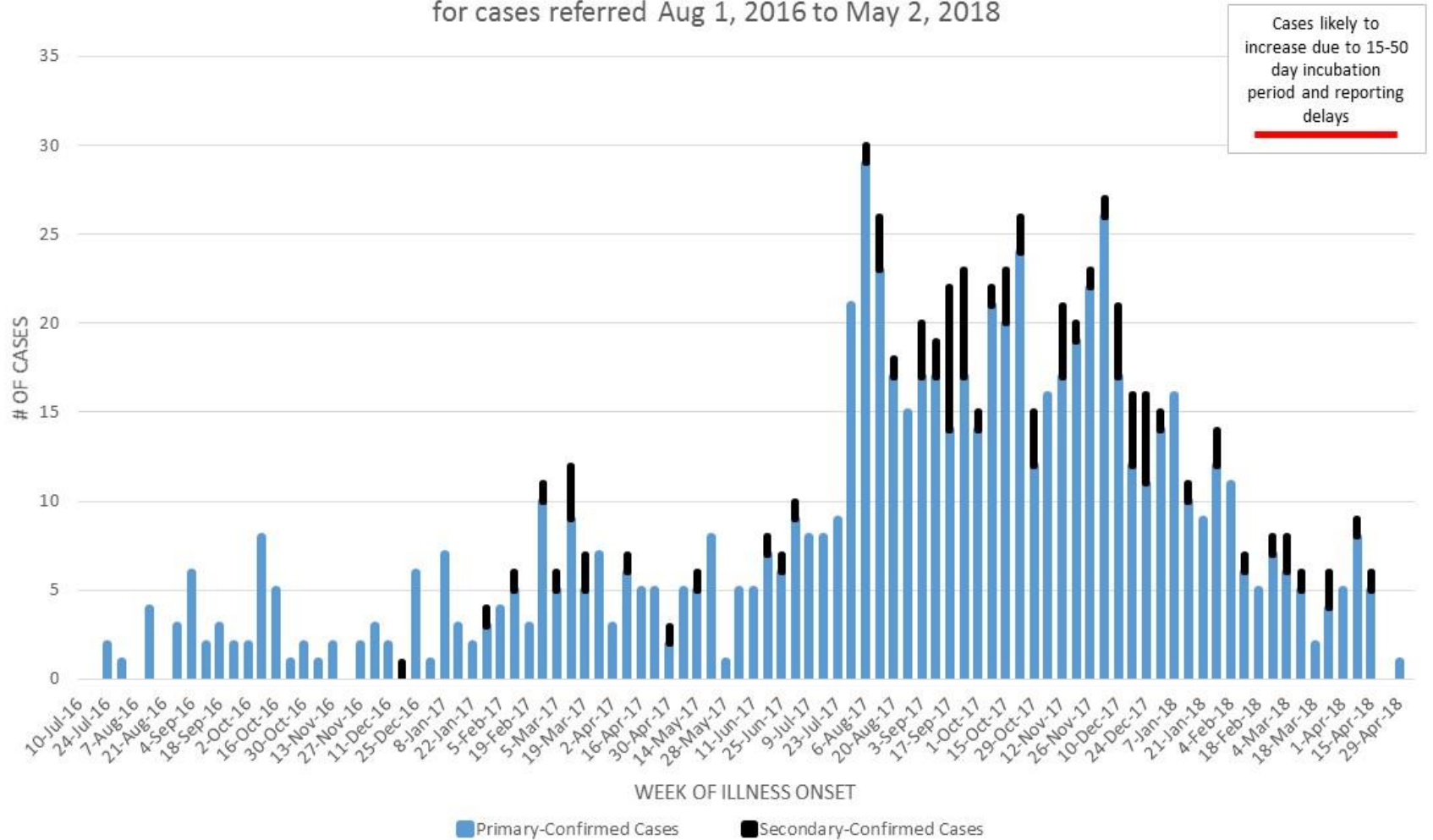
Epidemiologic Summary

Counts & Demographics

Epi Summary for Hepatitis A Cases in Michigan Reported Aug 1, 2016 – May 2, 2018

- 828 Total Cases
- 537 (64.8%) Male
- 665 (80.3%) Hospitalized
- Age range, <1–90 years
- Median age, 40 years
- 26 (3.1%) Deaths

Confirmed Hepatitis A Case Onset by Week for the Michigan Outbreak for cases referred Aug 1, 2016 to May 2, 2018



*If illness onset was not identified first lab collection date was used in place

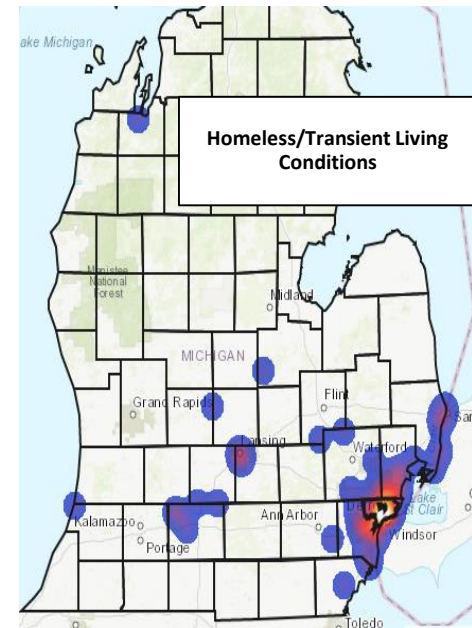
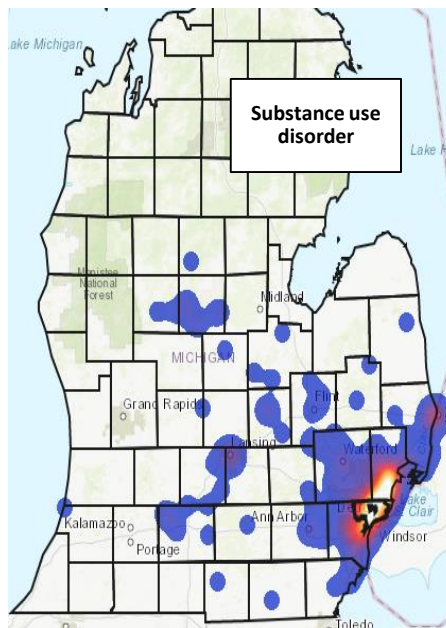
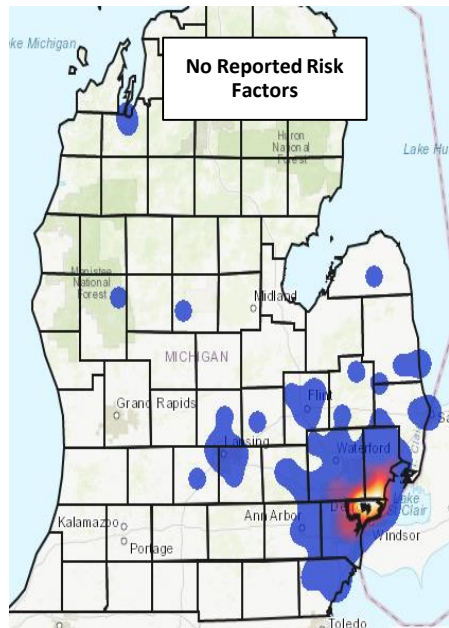
Epidemiologic Summary

Analysis of risk factor cases

Confirmed Hepatitis A by Risk Factors Reported August 1, 2016 to May 2, 2018

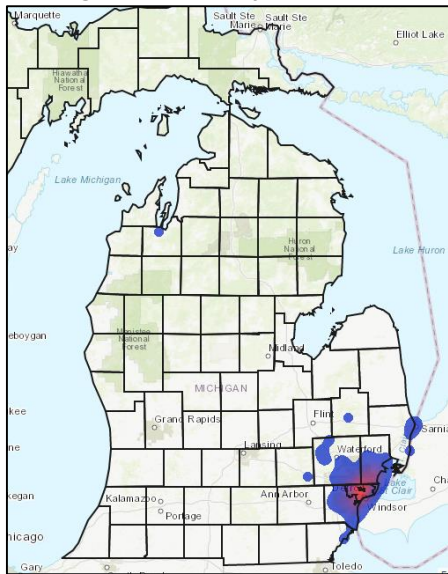
- 50.2% Documented Substance Use Disorder (374)
 - 8% Injection
 - 20% Non-Injection (most report marijuana use)
 - 20% Both (Injection & non-injection)
- 26.4% Coinfection with Hepatitis C (197)
- 14.6% Men Who Have Sex with Men (73)
- 13.4% Homeless or Transient Living (100)
- 7.7% Recently Incarcerated (57)
- 4.7% Food Worker (35)
- 3.0% Healthcare Worker (22)
- 2.7% Coinfection with Hepatitis B (20)

Relative Case Distribution by Risk Factor Status Aug. 2016 – Feb. 2018

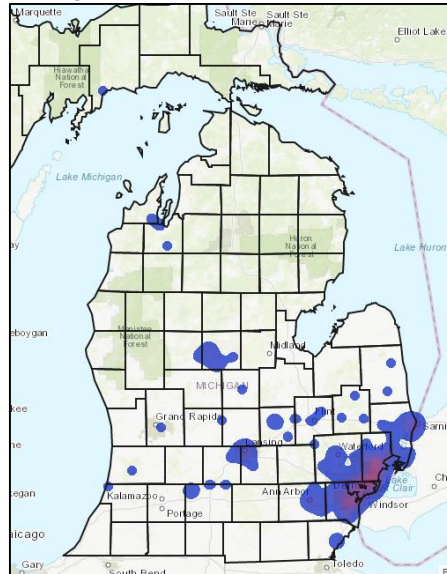


Temporal and Geographic Distribution of Outbreak Cases Patients with High Risk Characteristics

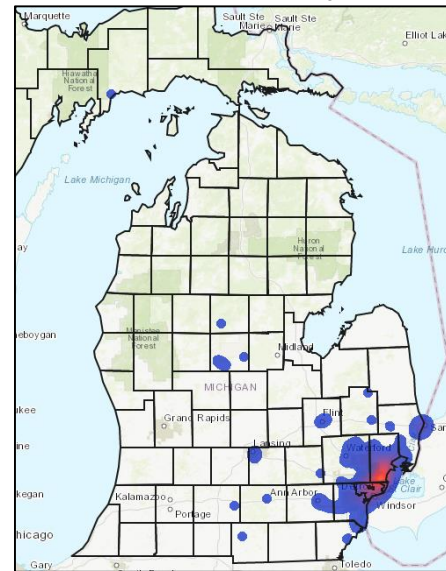
August 2016 – July 2017



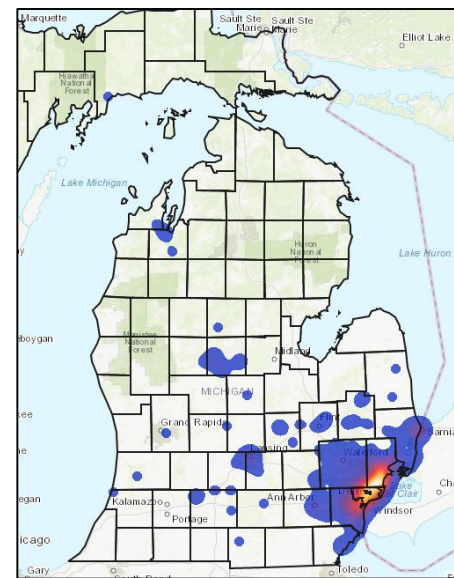
August 2017 – October 2017



November 2017 – January 2018



August 2016 – January 2018



Data source: Michigan Disease Surveillance System, MDHHS.

TRENDS IN THE NUMBER OF CASES

828

Cases

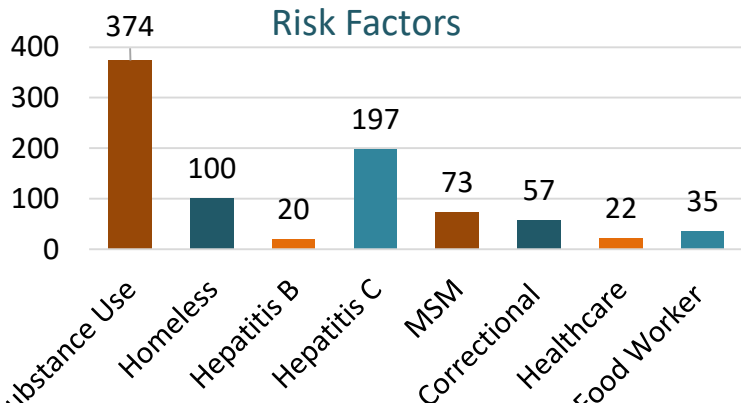
665

Hospitalized

26

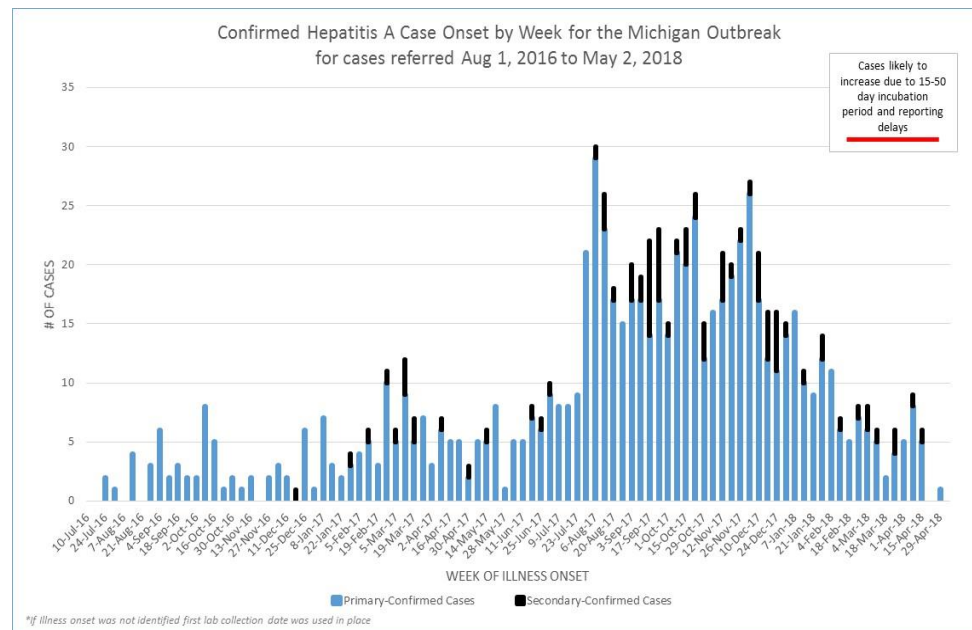
Deaths

Risk Factors



	February 2011-2016	February 2018	March 2011-2016	March 2018	April 2011-2016	April 2018
Min Cases	2	34	3	25	2	21
Max Cases	13		9		11	
Average # cases	7		6		6	

Source: MDSS



Guidelines for Management of Hepatitis A, Michigan Outbreak, 2018

PUBLIC HEALTH OUTBREAK RESPONSE

MDHHS and local public health officials are working to:

- Begin case investigation within 12 h after reported to public health
- Provide guidance and data to healthcare community
- Educate the public about hepatitis A and prevention
- Encourage community agencies and healthcare providers to immunize clients with risk factors for hepatitis A
- Increase availability of vaccine and conduct vaccination clinics
- **Increase vaccinations!**

Hepatitis A Vaccination for Outbreak Control, MI Outbreak

- Vaccination is the cornerstone of control of community outbreaks
- Post-exposure prophylaxis alone may not effectively control outbreaks
- Targeted vaccination to the groups at highest risk are the best way to control disease spread
- Primary prevention with adequate vaccination of at-risk groups is preferable

Hepatitis A Vaccination for Outbreak Control, MI Outbreak

- Vaccination in EDs was a major success in San Diego
- Screening tools are available on our website and from your peers
- Screen for insurance status and risk group
 - Public vaccine available for Medicare, Medicaid, and uninsured
 - Risk groups: sub, homeless, liver disease, MSM, recent incarceration
- Public doses must be registered in MCIR (MAVP)
- Not required to look up the individual in MCIR before vaccinating

Vaccination and PEP, MI Outbreak

Box A: Identifying PEP for patient based on age (years) and health status

Age	<1	1-40	41-59*	60-74*	75+
Healthy	IG	Vaccine Preferred	IG; vaccine if IG is in short supply	IG; vaccine if IG is in short supply	IG
Other (Box B)	IG	IG	IG	IG	IG
Highest Risk (Box C)	Consider vaccine and IG for possible longer-term protection				

*When IG is unavailable or in short supply, single-antigen HAV vaccine may be used for PEP in healthy people 41-74 years of age. To read more about hepatitis A vaccine for PEP in this age group, please see:

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC4643264/>.

Box B: People who are preferred to receive IG for PEP

- Those less than 12 months of age
- Those aged 41 through 74 years (*Vaccine can be used if IG is not available)
- Those who are immunocompromised, including persons:
 - With HIV/AIDS
 - Undergoing hemodialysis
 - Who have received solid organ, bone marrow or stem cell transplants
 - Receiving high dose steroids (>2mg/kg/day)
 - Receiving chemotherapy, immunomodulators and/or biologic medications, (mercaptopurine, methotrexate, infliximab, adalimumab, etanercept, tacrolimus, mycophenolate, etc.)
 - Persons who are otherwise less capable of developing a normal response to immunization
- Those who have chronic liver disease or other chronic medical conditions
- Those whom vaccine is contraindicated

Box C: People with High Risk Indications who should be considered for receiving IG AND hepatitis A vaccine for PEP^{2,3}

- Pregnant women
- Persons with chronic liver disease
- Persons who are immunocompromised, including persons:
 - With HIV/AIDS
 - Undergoing hemodialysis
 - Who have received solid organ, bone marrow or stem cell transplants
 - Receiving high dose steroids (>2mg/kg/day)
 - Receiving chemotherapy, immunomodulators and/or biologic medications, (mercaptopurine, methotrexate, infliximab, adalimumab, etanercept, tacrolimus, mycophenolate, etc.)
 - Persons who are otherwise less capable of developing a normal response to immunization

NOTES:

¹The efficacy of combined HAV/HBV (Twinrix®) vaccine for post-exposure prophylaxis (PEP) has not been evaluated so it is not recommended for PEP.

²Guidance was provided by a CDC subject matter expert with the Division of Viral Hepatitis on situations when IG and hepatitis A vaccine should be administered at the same time.

³If hepatitis A vaccine and IG are both considered then they may be administered simultaneously but at separate anatomic injection site.

MI OUTBREAK RISK FACTORS

Are you at risk for Hepatitis A?

People who are at high risk include:

- Men who have sex with men (MSM)
- People who use illegal drugs
- People currently homeless or in transient living
- People recently in jail or prison
- People with underlying liver disease*

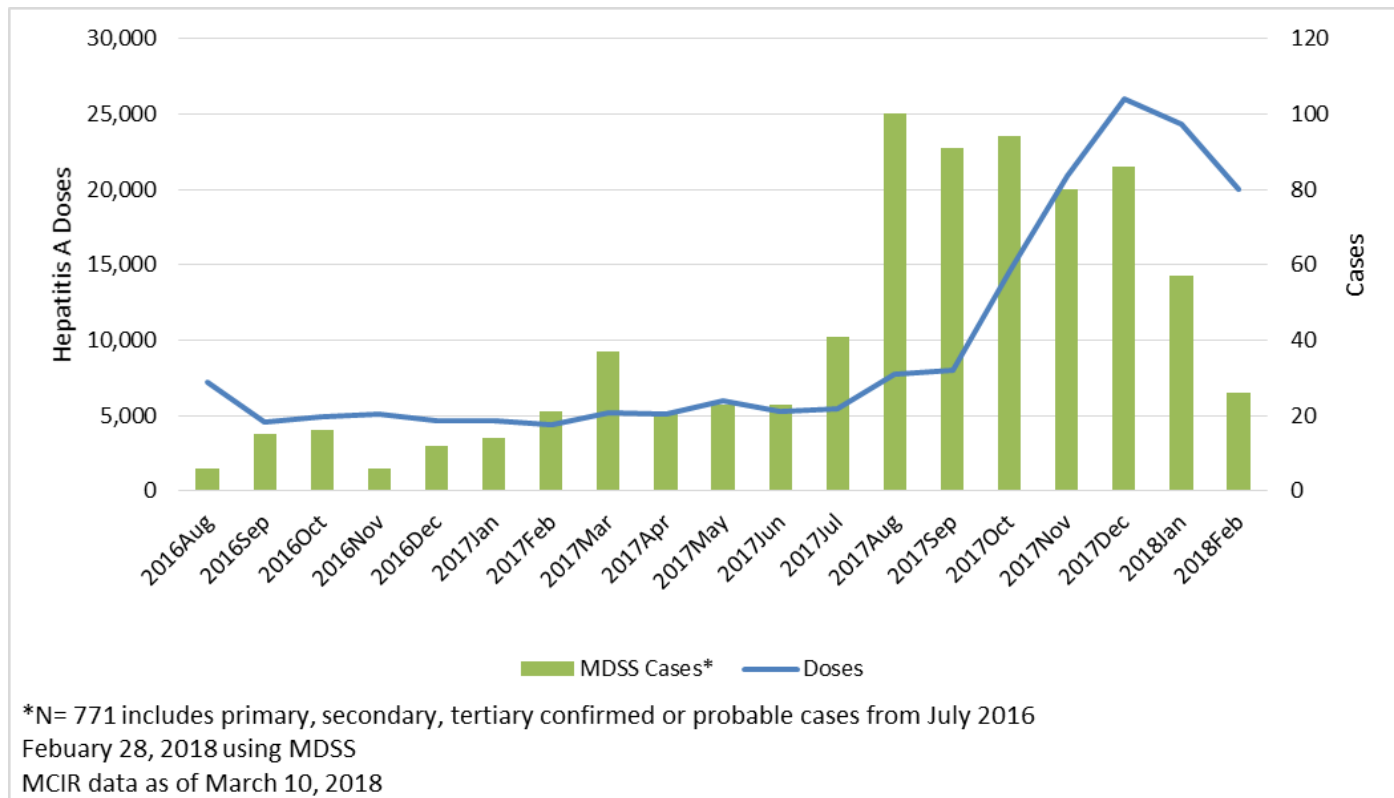
*Note: people with underlying liver disease (e.g., cirrhosis, hepatitis B, or hepatitis C) are at increased risk of having poor outcomes if they are infected with hepatitis A.

Ask your doctor about vaccination if you are at high risk.



The best way to protect against hepatitis A is to get the hepatitis A vaccine.

HAV cases vs. HAV doses administered to adults by month, August 2016 - February 2018



Getting the Vaccine out: Partners – A Selection

State

MDHHS Public Health Administration
Bureau of Epidemiology and Population Health
Bureau of Family Health Services
Bureau of Laboratories
Bureau of EMS, Trauma, and Preparedness
Bureau of Community-based Services
Bureau of Health and Wellness
Local Health Services
Medical Services
MI Volunteer Registry
External Affairs and Communications
Legislative and Constituent Services

Michigan Department of Corrections
Michigan Department of Agriculture and Rural Development

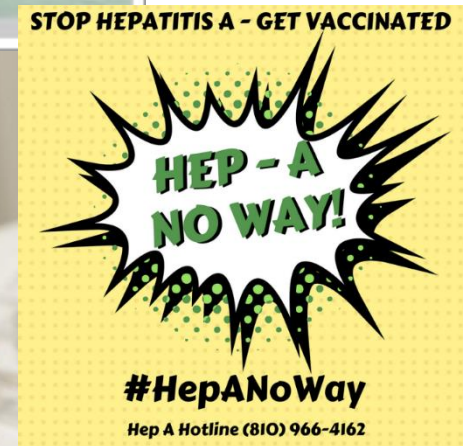
Professional

Michigan Health & Hospital Association
Michigan State Medical Society
Michigan Osteopathic Association
Michigan Association for Local Public Health
Michigan Primary Care Association
Michigan College of Emergency Physicians
Michigan Association of Community Mental Health Boards
Michigan Association of Family Physicians
Visiting Nurses Association

Partners – A selection (cont'd)

Local & Regional

Local health departments, Clinics
County Jails, Correct Care Solutions, and drug courts
Regional Healthcare Coalitions
Salvation Army Rehabilitation
Street Medicine Detroit
Neighborhood Service Organization - Tuminai Clinic
Detroit Recovery Project
Capuchin Soup Kitchen
Mariners Inn
Samaritas House
St. John Community Center
Detroit Rescue Mission
Naomi's Nest
Elmhurst Home
Self-Help Addiction Rehabilitation (SHAR)
Community Health Awareness Group (CHAG)
Sacred Heart Rehabilitation Centers
Community Programs, Inc.
Turning Point
Meridian



Waning Hep A Immunity in HIV, Michigan 2018

Possible loss of hepatitis A virus (HAV)
seroprotection noted among people
living with HIV — Michigan, 2018

Potential Waning Hep A Immunity in Patients Living with HIV

- Healthcare providers have reported that people living with HIV who were previously vaccinated against hepatitis A or had positive total HAV antibody testing may be susceptible and at risk for acquiring hepatitis A virus infection

Potential Waning Hep A Immunity in Patients Living with HIV

- 2 hepatitis A cases had positive total hepatitis A virus (HAV) antibody test results upon entry into care for HIV
 - They were not offered HepA vaccination previously because of presumed immunity
 - These providers have instituted re-screening patients who have not had total HAV antibody testing in the past 5 years
- Additional patients have been identified who have seroreverted from positive total HAV antibody status to negative, including those with history of HepA vaccination.

Potential Waning Hep A Immunity in Patients Living with HIV

- Though inconclusive, these early findings are concerning for loss of seroprotection in PLWH who may be susceptible and at risk of acquiring HAV infection.
- Total HAV antibody status should be updated if testing has not been performed during the previous 5 years for patients at risk during this outbreak
 - MSM, illicit substance use, homelessness or in transient living conditions, recent incarceration, and underlying liver disease including hepatitis B or C
- If total HAV Ab testing is negative, regardless of previous vaccination history, MDHHS recommends:
 - the patient completes the monovalent HepA vaccine series
 - the provider documents a post-vaccination response at least 4 weeks after the 2nd dose

Clinician Resources- Hep A and HIV

- Clinical experts at Henry Ford Hospital are available through the HIV Consultation Program for hepatitis A questions related to HIV patients.
- Non-urgent questions can be submitted at www.henryford.org/HIVconsult, and will be responded to in 24 to 48 hours.
- For urgent questions, health care professionals should contact the 24-hour consultation line by calling 313-575-0332.

Hepatitis A Outbreak Website

Hepatitis A Southeast Michigan Outbreak

Public health officials and the Michigan Department of Health and Human Services (MDHHS) are continuing to see an elevated number of hepatitis A cases in Southeast Michigan.

Since the beginning of the outbreak in August 2016, public health response has included increased healthcare awareness efforts, public notification and education, and outreach with vaccination clinics for high-risk populations. No common sources of food, beverages, or drugs have been identified as a potential source of infection. Transmission appears to be through direct person-to-person spread and illicit drug use. Those with history of injection and non-injection drug use, homelessness or transient housing, and incarceration are thought to be at greater risk in this outbreak setting. Notably, this outbreak has had a high hospitalization rate.



Southeast Michigan Hepatitis A Outbreak Cases and Deaths as of December 13, 2017*

*Table will be updated weekly by 4:00pm each Friday

Cases	Hospitalizations	Deaths
610	501 (82.1%)	20 (3.3%)

Please note: Table does not include all reported hepatitis A cases in the SE MI outbreak region; only those cases that are identified as outbreak-related. More descriptive data on the current outbreak can be found within the [Comprehensive Summary](#). Data are provisional and subject to change.

In support of efforts, MDHHS has a website for the hepatitis A outbreak that has important and timely information, available at:

➤ www.mi.gov/hepatitisAoutbreak

- The website contains a brief case count, hospitalized cases, and deaths for an at-a-glance review that is updated each Friday.
 - Confirmed cases are also listed out by jurisdiction.
 - A Comprehensive Summary with case demographics and risk profiles is also available
- A full listing of communication documents and educational materials available for download.

Printed Materials



[Brochure – Help stop the spread of hepatitis A in Michigan communities](#)
(updated 2/8/2018)



[Poster – Protect Yourself from Hepatitis](#)
(updated 11/2017)

Available to order in the
Clearinghouse at no cost!
(Brochure on backorder, being re-printed)

<http://www.hpclearinghouse.org/>
Click “Enter Here to Place Your Order”
Click “Immunizations”

Item Numbers:
IM160 – Poster
IM161 – Brochure

- Brochures translated in Arabic, Spanish, Chinese, and Bengali available at:
www.mi.gov/hepatitisAoutbreak

Outreach Materials

Help Stop the Spread of Hepatitis A

Hepatitis A is spreading in Southeast Michigan. Hepatitis A is a serious liver disease that can easily be passed from person to person. Anyone can get hepatitis A. It spreads when people do not wash their hands before making food and drinks. It also spreads through close contact with people who have hepatitis A. Washing your hands with soap and warm water after using the bathroom, changing a diaper, or before making food and drinks can help stop the spread of hepatitis A.

► [Flyer – Help Stop the Spread of Hepatitis A](#)

Hepatitis A is Spreading

among MSM and persons who inject drugs in Southeast Michigan

HEPATITIS A IS A SERIOUS LIVER DISEASE CAUSED BY A VIRUS THAT IS FOUND IN THE FECES (POOP) OF INFECTED PEOPLE. IT CAN EASILY SPREAD FROM PERSON-TO-PERSON AND CAUSE LIVER DISEASE LASTING A FEW WEEKS TO A SERIOUS ILLNESS LASTING MANY MONTHS. IN SOME CASES, IT CAN BE DEADLY.

Protect yourself against this outbreak by getting your first dose of hep A vaccine.

Wash Your Hands

1. **Wet** your hands with clean, running warm water and apply soap.
2. **Lather** your hands by rubbing them together with the soap. Be sure to lather the backs of your hands, between your fingers, and under your nails.
3. **Scrub** your hands for at least 20 seconds.
4. **Rinse** your hands under clean, running water.
5. **Dry** your hands with a clean towel or air dry.

*Alcohol based hand sanitizers are not effective against the hepatitis A virus.

The best way to protect against hepatitis A is to get the hepatitis A vaccine. Talk to your healthcare provider about getting vaccinated. If you need help paying for the vaccine, your local health department may have vaccine available for little or no cost.

For more information visit:
www.mi.gov/hepatitisAoutbreak
or call 1-800-872-2437 M-F 9am-5pm

Protect yourself. Get vaccinated today.

MDHHS
Michigan Department of Health & Human Services

► [Flyer – Hepatitis A is in Michigan communities](#)

Hepatitis A is in Michigan communities.



Hepatitis A is a liver disease caused by the hepatitis A virus (HAV). Hepatitis A is spread through contaminated food or water and close contact with persons who are infected. Hepatitis A can affect anyone. Frequent hand washing with soap and warm water after using the bathroom, changing a diaper, or before preparing food can help prevent the spread of Hepatitis A.

The best way to protect against hepatitis A is to get the hepatitis A vaccine. Talk to your health care provider to get the two doses you need for protection. Need help paying for vaccines? Your local health department or your federally qualified health center may have hepatitis A vaccine available for little cost.

▼

Stop the spread. Get vaccinated today.

www.michigan.gov/immunize

MDHHS
Michigan Department of Health & Human Services
1000 W. Wacker Drive, Lansing, MI 48906

Questions