INFECTIOUS DISEASES IN THE MOVIES: FACT OR FICTION

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Technical Assistance: Gordon Palmer UNC Medical Illustration & Photography

LECTURE GOALS

- □ Provide entertainment (learning medicine should be fun)
- □ Provide trivia for your next cocktail party or Jeopardy appearance
- Develop an appreciation for classic movies
- Place infectious diseases in historical perspective

DISCLOSURES

- □ No honoraria or grants from film companies
- No mention of off label use of FDA approved drugs (in fact no mention of drugs at all)
- □ As an intern I took care of Marion Morrison

DISCLOSURES

- No honoraria or grants from film companies
- No mention of off label use of FDA approved drugs (in fact no mention of drugs at all)
- As an intern I took care of Marion Morrison (i.e., John Wayne or "The Duke")



TRIVIA QUIZ: QUESTIONS 1 & 2

- $\hfill \square$ Most common infectious disease in US presidents
 - A. Tuberculosis
 - B. Malaria
 - C. Syphilis
 - D. Rheumatic fever
- $\hfill\Box$ Number of presidents with the disease
 - A. 3
 - B. 7
 - C. 11
 - D. 13

TRIVIA QUIZ: ANSWERS

- $\hfill \square$ Most common infectious disease in US presidents
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 - D. 13

MALARIA, US



© Elsevier 2004. Infectious Diseases 2e - www.idreference.com

INFECTIOUS DISEASES IN THE MOVIES

Historical movies-overcoming adversity

- □ Miracle Worker, 1962: scarlet fever
- □ Tombstone, 1993: Tuberculosis

Period movies - overcoming adversity

- □ Five Pennies, 1959: Polio
- □ Philadelphia, 1993: AIDS
- □ Finding Neverland, 2004: Tuberculosis

Tear jerkers

- □ Little Women, 1949: Scarlet fever
- □ Old Yeller, 1957: Rabies

Infectious diseases as natural hazards

- □ African Queen, 1951: Mosquitoes, leeches
- □ To Kill A Mocking Bird, 1962: Rabies
- □ Motorcycle Diaries, 2004: Leprosy

Scary movies

- □ War Of The Worlds, 1953: "Common" bacteria
- □ Outbreak, 1995: Ebola
- □ Cujo, 1983: Rabies



NEW DISEASES: THE FACTS

- □ Mars is the most earth-like of the planets
- □ Many recent epidemics have been due to microbes jumping species lines
- □ We live in a sea of "organisms"
- □ Fortunately, our host defenses, in general, prevent us form acquiring infections from most environmental microbes (most infections come from endogenous flora)

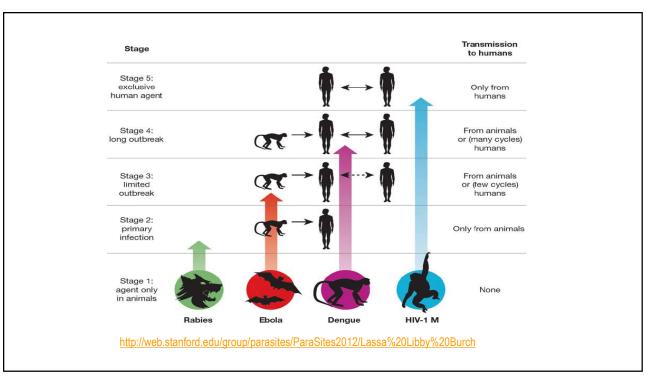
NEW DISEASES: FICTION

- □ No life on mars
- Most microbes that have jumped species lines and caused recent epidemics have been viruses (not bacteria)
- Host resistance varies; implausible that an entire species would be killed by a single microbe simultaneously

HISTORY OF PANDEMICS AND EMERGING DISEASES Death toll Type / Pre-human host DEATH TOLL Antonine Plague 165-180 Believed to be either smallpox or measles Japanese smallpox epidemic 200M 735-737 Variola major virus 1M Plague of Justinian 541-542 Yersinia pestis bacteria / Rats, fleas Black Death 1347-1351 Yersinia pestis bacteria / Rats, fleas 200M New World Smallpox Outbreak 1520 – onwards Variola major virus 56M Great Plague of London 1665 Yersinia pestis bacteria / Rats, fleas Cholera Pandemics 1-6 1817-1923 Third Plague Yellow Fever Virus / Mosquitoes 100,000-150,000 (U 1918-1919 1957-1958 1.1M Asian Flu 1968-1970 Hong Kong Flu 1981-present HIV/AIDS Virus / Chimpanzees 2009-2010 H1N1 virus / Pigs 200,000 Swine Flu SARS 2002-2003 Coronavirus / Bats, Civets Ebola 2014-2016 Ebolavirus / Wild animals 11,000 MERS 2015-Coronavirus / Bats, camels 850 Present COVID-19 2019-Present 14,500 (as of Mar 2 2020) pangolins) Note: Many of the death toll numbers listed above are best estimates based on available research. Some, such as the i of Justinian, are <u>subject to debate</u> based on new evidence.

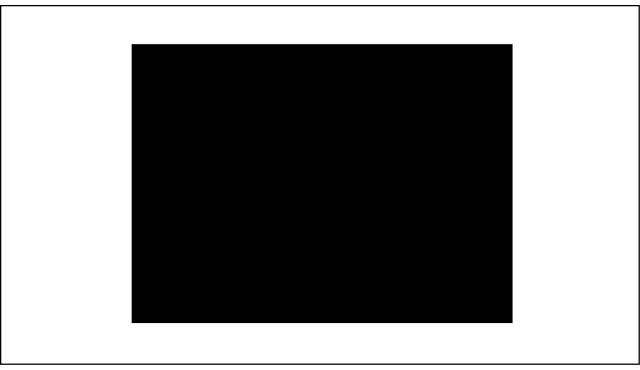
EMERGING DISEASES IN THE US

DISEASE (source)	CASES	OUTCOME	YEAR
West Nile virus (Israel)	Thousands	Endemic (US)	1999
SARS (China)	8096 (8 US, 1 UNC)	Controlled	2003
Monkeypox (Africa)	71	Controlled	2003
Novel flu, H1N1 (Mexico)	Thousands	Endemic (Worldwide)	2009
MERS-CoV (Arabian Peninsula)	Hundreds	Epidemic (Arabian area)	2014
Enterovirus D68	Hundreds (13 UNC)	Epidemic (US)	2014
Ebola	Thousands (1 US)	Epidemic (West Africa)	2014-15
SARS-CoV-2	Millions	Endemic (Worldwide)	2019-present



BASIC CONCEPTS IN DISEASE EMERGENCE

- □ Emergence of infectious diseases is complex
- □ Infectious diseases are dynamic
- Most new infections are not caused by genuinely new pathogens
- Agents involved in new and reemergent infections cross taxonomic lines
- □ The concept of the microbe as *the* cause of disease is inadequate and incomplete
- Human activities are the most potent factors driving disease emergence
- □ Social, economic, political, climatic, technologic, and environmental factors shape disease patterns and influence emergence
- □ Understanding and responding to disease emergence require a global prospective, conceptually and geographically
- ☐ The current global situation favors disease emergence



EBOLA: THE FACTS

- □ Multiple outbreaks described in Africa; first outbreak in Zaire 1967
- □ Acquisition from monkeys
- □ High mortality
- □ Person-to-person transmission; healthcare personnel at high risk
- □ Therapy has advanced from supportive to monoclonal antibodies approved in 2020 (Inmazeb, Ebanga)
- □ Pre-exposure prophylaxis now also available with rVSV-ZIBOV (Ervoba), approved by FDA in 2019

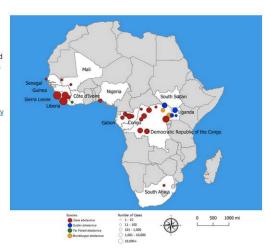
EBOLA: FICTION

- □ The US doesn't destroy villages suffering an outbreak of Ebola
- □ The reservoir is probably bats
- □ Pandemic unlikely; high mortality and short incubation period

Emergence of Ebola in Humans

Viral and epidemiologic data suggest that Ebola virus existed long before these recorded outbreaks occurred. Factors like population growth, encroachment into forested areas, and direct interaction with wildlife (such as bushmeat consumption) may have contributed to the spread of the Ebola virus.

Since its discovery in 1976, the <u>majority</u> of cases and outbreaks of Ebola Virus
Disease have occurred in <u>Africa</u>. The
2014-2016 Ebola outbreak in West
Africa began in a rural setting of
southeastern Guinea, spread to urban
areas and across borders within
weeks, and became a global epidemic
within months.



https://www.cdc.gov/vhf/ebol a/history/summaries.html

Chronology of Previous Ebola Virus Disease Outbreaks, WHO Year 2021 Cases Ongoing Uganda 44% 2021 2003 (Nov-Dec) 2003 (Jan-Apr) 2001-2002 Uganda South Africa (ex-Gabon) 2000 53% 1996 100% 2015 2014 2014 Zaire Zaire Zaire Zaire Zaire Zaire 1996 (Jan-Apr) Gabon 2014 2014 2014 Côte d'Ivoire Democratic Republic of the 100% Republic of the Congo Democratic Republic of the Congo Uganda Uganda Democratic Republic of the Congo Sudan Sudan

EBOLA VIRUS DISEASE, KEY FACTS, WHO

- □ Ebola virus disease (EVD), formerly known as Ebola haemorrhagic fever, is a rare but severe, often fatal illness in humans.
- The virus is transmitted to people from wild animals and spreads in the human population through human-to-human transmission.
- □ The average EVD case fatality rate is around 50%. Case fatality rates have varied from 25% to 90% in past outbreaks.
- Community engagement is key to successfully controlling outbreaks.
- Good outbreak control relies on applying a package of interventions, namely case management, infection prevention and control practices, surveillance and contact tracing, a good laboratory service, safe and dignified burials and social mobilization.
- Use Vaccines to protect against Ebola have been developed and have been used to help control the spread of Ebola outbreaks in Guinea and in the Democratic Republic of the Congo (DRC).
- Early supportive care with rehydration, symptomatic treatment improves survival. Two monoclonal antibodies (Inmazeb and Ebanga) were approved for the treatment of Zaire ebolavirus (Ebolavirus) infection in adults and children by the US Food and Drug Administration in late 2020.
- Pregnant and breastfeeding women with Ebola should be offered early supportive care. Likewise vaccine prevention and experimental treatment should be offered under the same conditions as for non-pregnant population.

https://www.who.int/news-room/fact-sheets/detail/ebola-virus-disease

KEY CONSIDERATIONS IN ASSESSING AND MANAGING THE THREAT OF AN EMERGING INFECTIOUS DISEASE

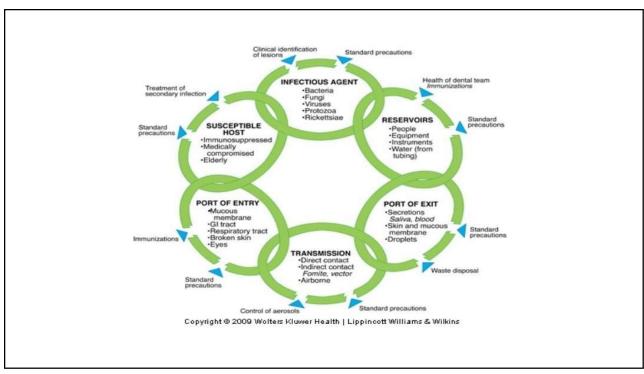
- Pathogen
 - Taxonomy (provides clues regarding transmission routes, environmental stability, germicide susceptibility)
 - Hosts
- Epidemiology
 - Locations of endemicity (i.e., locations in the world where sources or reservoirs reside)
 - Incubation period
 - Transmission routes
 - Infectivity (i.e., communicability)
 - Duration of infectivity

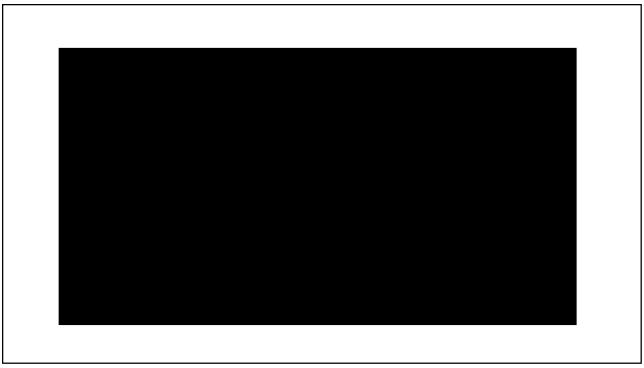
- Clinical
 - Symptoms
 - Signs
 - Risk factors for acquisition of infection
 - Morbidity
 - Mortality
 - Risk factors for morbidity and mortality
 - Diagnostic methods (sensitivity, specificity, biosafety)
 - Therapy (availability, efficacy, safety)

KEY CONSIDERATIONS IN ASSESSING AND MANAGING THE THREAT OF AN EMERGING INFECTIOUS DISEASE

- Infection Prevention
 - Environmental survival
 - Germicide susceptibility
 - Isolation recommendations
 - Recommended personal protective equipment
 - Pre-exposure prophylaxis (availability, efficacy, safety)
 - Postexposure prophylaxis (availability, efficacy, safety)
 - Recommended biosafety level in the laboratory
 - Recommended waste disposal (liquids and solids

- Managing a pandemic
 - Sensitive and specific (ideally rapid) diagnostic test
 - Early identification of patients
 - Protecting our healthcare personnel (PPE, donning, doffing)
 - Sufficient staff, inpatient/ICU beds, ventilators
 - Managing shortages







RABIES: EPIDEMIOLOGY

- □ Agent: Rhabdoviridae (genus Lyssavirus)
- □ Reservoir: Small mammals, bats
 - Bat species include silver-haired bats and eastern pipistrelles
- □ Transmission
 - Animal bite or scratch (contact with infectious saliva)
 - Airborne: Spelunking (rare) or laboratory generated aerosol
 - latrogenic: Corneal transplants (8 cases), transplanted organs (2 outbreaks) or improperly inactivated vaccines
- Deaths, US
 - From 1960 to 2018, 127 human rabies cases were reported in the United States, with roughly a quarter resulting from dog bites received during international travel. Of the infections acquired in the United States, 70% were attributed to bat exposures.
- □ Prevention (or post-exposure therapy)
 - Rabies vaccine + RIG

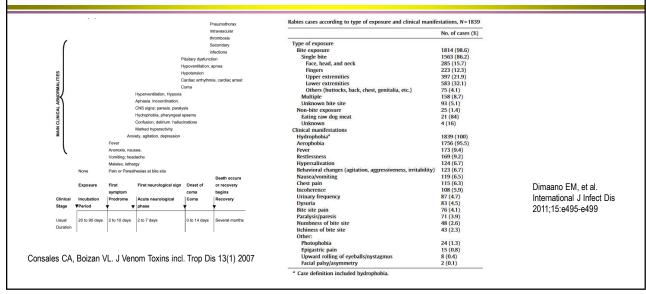
Estimated Incubation Period (days) per Bite Site

Bite site	<7 (%)	8-14 (%)	15-21 (%)	22-30 (%)	31-90 (%)	91-365 (%)	>365 (%)
Face, head, neck	5 (0.3)	32 (1.8)	49 (2.8)	57 (3.3)	84 (4.8)	46 (2.6)	12 (0.7)
Fingers	0	0	13 (0.7)	21 (1.2)	44 (2.5)	106 (6.1)	39 (2.2)
Upper extremities	0	0	3 (0.2)	8 (0.5)	76 (4.4)	225 (12.9)	85 (4.9)
Lower extremities	0	0	0	5 (0.3)	58 (3.3)	369 (21.2)	151 (8.7)
Other sites (buttocks, back, chest, genitalia)	0	0	1 (0.1)	6 (0.3)	23 (1.3)	31 (1.8)	14 (0.8)
Multiple bite	2 (0.1)	5 (0.3)	16 (0.9)	24 (1.4)	69 (4.0)	34 (2.0)	8 (0.5)
Non-bite (eating raw dog meat)	0	0	0	3 (0.2)	7 (0.4)	11 (0.6)	0
Total cases ^b	7 (0.4)	38 (2.1)	92 (5.0)	155 (8.4)	498 (27.1)	785 (42.8)	251 (13.7

Time of exposure to the time of manifest signs and symptoms; eight cases were of 5 years incubation (0.4%) and one case was of 27 years incubation (0.1%).
 Not all bite locations were recorded, thus n=1835 for total cases and n=1742 for all cases for which a bite location was recorded.

Dimaano EM, et al. International J Infect Dis 2011;15:e495-e499

RABIES, FREQUENCY ABND TIME COURSE OF SYMPTOMS



Rabies virus is adapted to its reservoir host and different variants exists in the United States, such as racoon variant, bat variant, fox variant, and skunk variant. Although cross-species transmission of rabies virus variants does occur (for example, infection of dogs with raccoon rabies variant), rabies virus variants are primarily transmitted within the species they are adapted to, such as the raccoon variant primarily being transmitted between raccoons. Rabies virus variants associated with the major mesocarnivore species (such as raccoons, skunks, foxes, and mongooses) are distributed in distinct geographic regions.

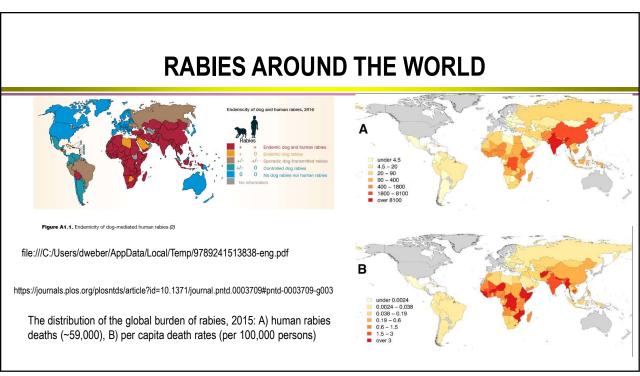


https://www.cdc.gov/rabies/location/usa/surveillance/wild_animals.html

Animal Rabies, US

Between 2013-2017, bats with rabies were found in every state except for Hawaii. Skunks with rabies have been found in parts of California, the Midwest, Texas, Kentucky, Virginia, North Carolina, and Tennessee. Raccoons with rabies have been found in the South and Eastern states. Foxes with rabies have been found in Alaska, Arizona, and New Mexico. Foxes and skunks with rabies have been found in Arizona, New Mexico, and Texas. Mongoose with rabies have been found in Puerto

HUMAN AND ANIMAL RABIES BY YEAR, NC Human Rabies Reported Cases North Carolina, 1929 - 2020 (n = 26) Confirmed Rabid Mammals by Year, North Carolina, 1991 - 2020 (n = 12,746) 1000 900 First (N.C.) Raccoon Variant Case Jones County, 2011 Raccoon Rabies; 1st Identified 1991 800 No. of Cases 700 600 500 400 Last reported Canine Variant Case Cherokee County, 1955 No. of Cases 300 200 100 *ERV = Eastern Raccoon Variant Source: NC State Laboratory of Public Health Updated Jan 2021 Year https://epi.dph.ncdhhs.gov/cd/rabies/figures.html#graphs



RABIES: EPIDEMIOLOGY AND PREVENTION, WHO

- □ Rabies is a vaccine-preventable viral disease which occurs in more than 150 countries and territories.
- Dogs are the main source of human rabies deaths, contributing up to 99% of all rabies transmissions to humans.
- Interrupting transmission is feasible through vaccination of dogs and prevention of dog bites.
- □ Infection causes tens of thousands of deaths every year, mainly in Asia and Africa.
- □ Globally rabies causes an estimated cost of US\$ 8.6 billion per year
- □ 40% of people bitten by suspect rabid animals are children under 15 years of age.
- □ Immediate, thorough wound washing with soap and water after contact with a suspect rabid animal is crucial and can save
- Engagement of multiple sectors and One Health collaboration including community education, awareness programes and vaccination campaigns are critical.

https://www.who.int/news-room/fact-sheets/detail/rabies

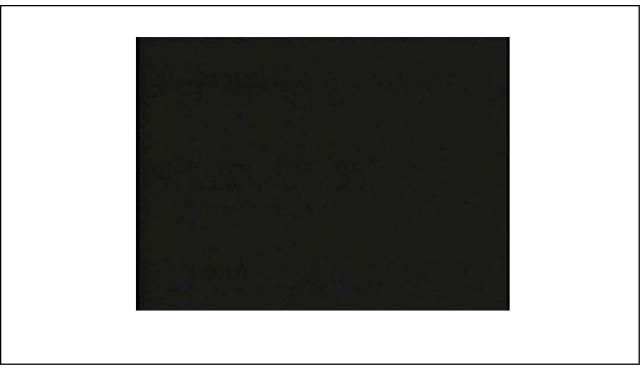


RABIES: THE FACTS

- □ Rabies endemic throughout the world
- Main source of human rabies in the US in the past was dogs
- □ Highly fatal to all mammals (only 6 human survivors)
- □ No treatment once symptoms appear
- □ In South America ruminants are a major source of human rabies (due to vampire bats)
- □ Animal saliva is source of infection (dead animals are indeed dangerous)

RABIES: FICTION

- □ All rabid animals are aggressive "furious rabies"
 - In 15-20% of cases the infected animal or human demonstrates the paralytic form ("dumb rabies")
- Most human cases in US were due to domestic or wild animal bites (most cases in US are now due to bats)
- Domestic animals rarely infected since dogs must be immunized in all states (cats also in most states including NC)



POLIO: THE FACTS

- □ Summer outbreaks were common
- □ Children most commonly infected
- □ Death commonly due to respiratory failure (therapy = support using a ventilator)
- □ Prior to vaccine, >20,000 cases per year of paralytic polio
 - The number of polio cases in the U.S. peaked at 57,879, resulting in 3,145 deaths.
- □ 1987 survey revealed 670,00 Americans living with sequelae of paralytic polio



Hospital room in the US, 1952

https://ourworldindata.org/polio

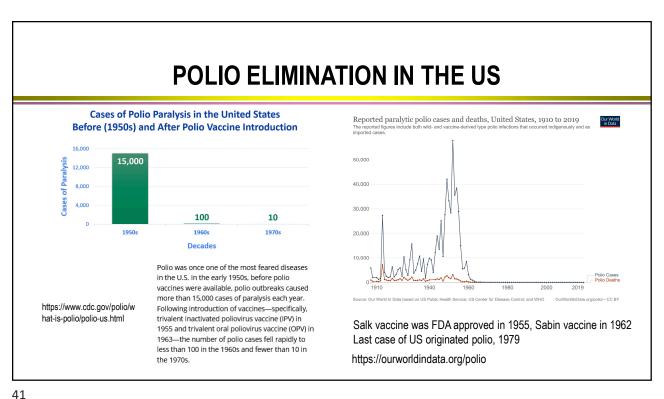
POLIO: FICTION

- □ Polio not transmitted by respiratory route; masks not necessary (fecal-oral transmission)
- □ Following acute polio, progressive improvement is the usual course (post-polio syndrome {PPS} affects ~25-40% of survivors, usually 15-40 years after the initial infection; many survivors experience a modest decline in function and muscle strength over many years)
 - Symptoms of PPS: Muscle weakness; mental and physical fatigue; joint pain
 - Some people with PPS have only minor symptoms, while others develop more visible muscle weakness and atrophy (a decrease in muscle size). PPS is rarely life-threatening, but the symptoms can make it difficult for an affected person to function independently.

INACTIVATED POLIO VACCINE (SALK) PROGRAM, 1955



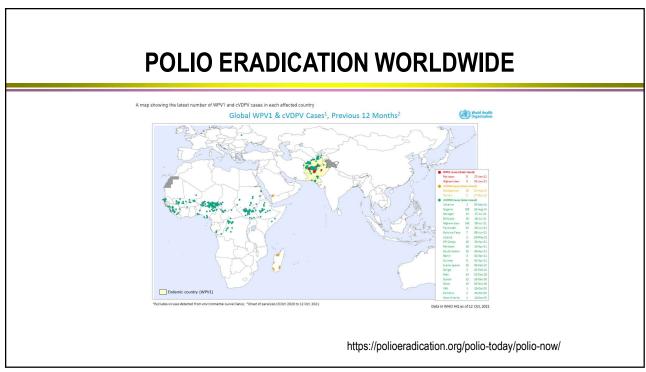




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POLIO ERADICATION

- Poliomyelitis (polio) is a highly infectious viral disease that largely affects children under 5 years of age. The virus is transmitted by person-to-person spread mainly through the fecal-oral route or, less frequently, by a common vehicle (e.g. contaminated water or food) and multiplies in the intestine, from where it can invade the nervous system and cause paralysis.
- In 1988, the World Health Assembly adopted a resolution for the worldwide eradication of polio, marking the launch of the Global Polio Eradication Initiative, spearheaded by national governments, WHO, Rotary International, the US CDC, UNICEF, and later joined by the Bill & Melinda Gates Foundation and Gavi, the Vaccine Alliance. Wild poliovirus cases have decreased by over 99% since 1988, from an estimated 350,000 cases in more than 125 endemic countries then to 175 reported cases in 2019.
- □ Of the 3 strains of wild poliovirus (type 1, type 2 and type 3), wild poliovirus type 2 was eradicated in 1999 and no case of wild poliovirus type 3 has been found since the last reported case in Nigeria in November 2012. Both strains have officially been certified as globally eradicated. As at 2020, wild poliovirus type 1 affects two countries: Pakistan and Afghanistan.
- The strategies for polio eradication work when they are fully implemented. This is clearly demonstrated by India's success in stopping polio in January 2011, in arguably the most technically challenging place, and polio-free certification of the entire WHO Southeast Asia Region in March 2014.



POLIO ERADICATION AND RESURGENCE

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- 2021, cVDPV2 was reported in 2021 in Ethiopia (2), Mali (1), Nigeria (23), Sierra Leone (1), and Tajikistan (1); 2022 cVDP2 reported in Israel, UK (positive wastewater led to vaccination of London children under 10) and NY (positive wastewater and 1 patient with paralytic polio)
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PARALYTIC POLIO, US

- Transmission: Fecal-oral (direct contact, indirect via contaminated water or food); droplet – infected people can shed virus for weeks from GI tract; virus can survive for weeks in the environment
- Long-term carrier states (>6 mo) may occur in immunocompromised persons
- Most cases of polio are asymptomatic but 25% develop flu-like symptoms: Pharyngitis, fever, fatigue, nausea, headache, abdominal pain – duration 2-5 days
- Meningitis = 1% to 5%; Paralysis = 0.5% to 0.05% (Between 2 and 10 out of 100 die)
- Post-polio syndrome: Development of new muscle pain, weakness, or paralysis as adults,15 to 40 years after recovery (25%-40% of patient with hx paralytic polio).
- Diagnosis: Detection in stool
- □ Vaccine: Only inactivated vaccine available in US (risk VAPP with oral vaccine, ½.4 million); routine childhood vaccination; effectiveness (3 doses) 99%-100%; >90% have antibody after 25 years

ummary

What is already known about this topic?

Sustained poliovirus transmission has been eliminated from the United States for approximately 40 years; vaccines are highly effective in preventing paralysis after exposure.

What is added by this report?

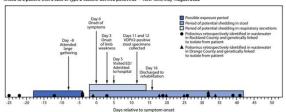
What he added by this report.

In June 2022, poliovirus was confirmed in an unvaccinated immunocompetent adult resident of New York hospitalized with flaccid lower limb weakness. Vaccine-derived poliovirus type 2 was isolated from the patient and identified from wastewater samples in two neighboring New York counties.

What are the implications for public health practice?

Unvaccinated persons in the United States remain at risk for paralytic poliomyelitis if they are exposed to either wild or vaccine-derived poliovirus; all persons in the United States should stay up to date on recommended poliovirus vaccination

FIGURE. Timeline of patient activities, potential poliovirus exposures, shedding, and poliovirus-positive wastewater* samples* genetically

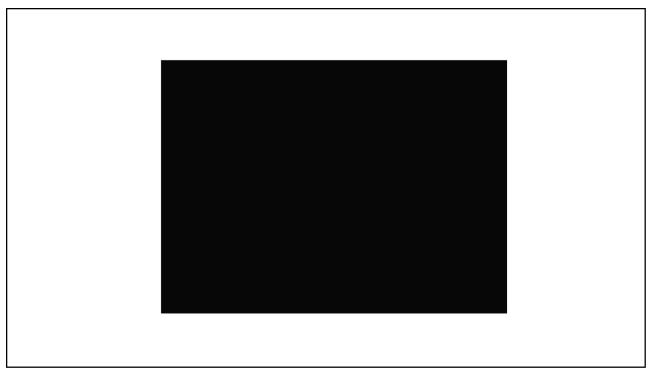


Abbreviations: ED -- emergency department VDP/2 -- type 2 vaccine- derived policivirus.

**Watestwater, abo eferred to as seeinge, includes water from household or building use (eg., tollets, showers, and sinks) that can contain human fecal waste a water from non-household sources (eg., rain and industrial use).

**More than one positive wastewater sample might have been collected on the same day in Rockland Country or Orange Country.

Link-Gelles R, et al. MMWR 2022;71:1065





FAMOUS PEOPLE WITH TUBERCULOSIS

□ *C. P. E. Bach: Musician

□ *Frederic Chopin: Musician

□ Noel Coward: Author

□ Ralph Waldo Emerson: Writer

Paul Erhlich: Physician

□ *Doc Holliday: Gunman

□ *Vivian Leigh: Actress

□ Franklin Pierce: President

☐ F. Scott Fitzgerald: Writer

□ *George Orwell: Writer

□ *Eleanor Roosevelt: Wife of president

□ *Henry David Thoreau: Writer

□ Leon Trotsky: Writer

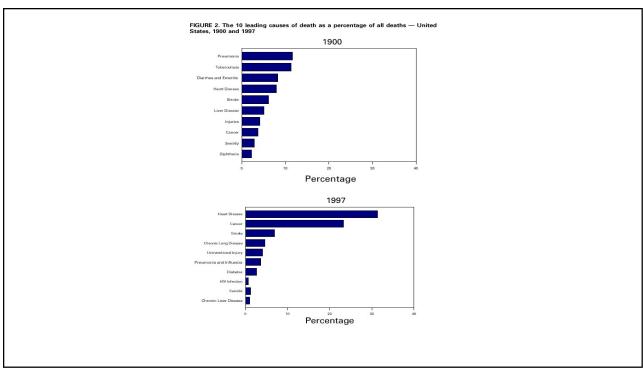
□ George Washington: President

□ *Walt Whitman: Writer

* Died of tuberculosis

DR. JOHN HENRY "DOC" HOLLIDAY

- □ 1851: Born in Griffin, Georgia, to a wealthy family
- □ 1870-72: Studied dentistry
- 1873: Found to have tuberculosis (possible nosocomial acquisition) and moved West after being told he had "3 months to live"
- □ 1873: Gave up dentistry due to his chronic cough
- □ 1875-1882: Killed >10 men
- □ 1882: Fought along the Earps at O.K. Corral
- □ 1887: Died of tuberculosis

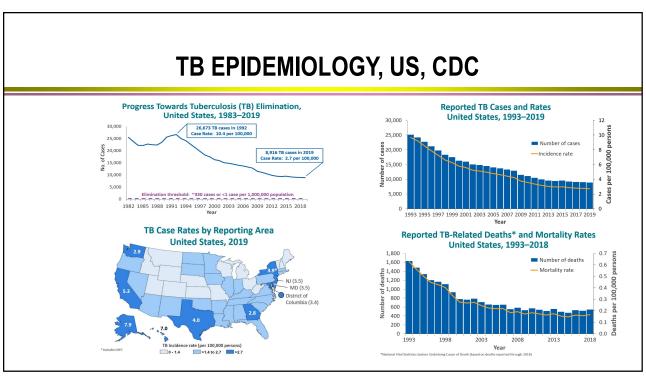


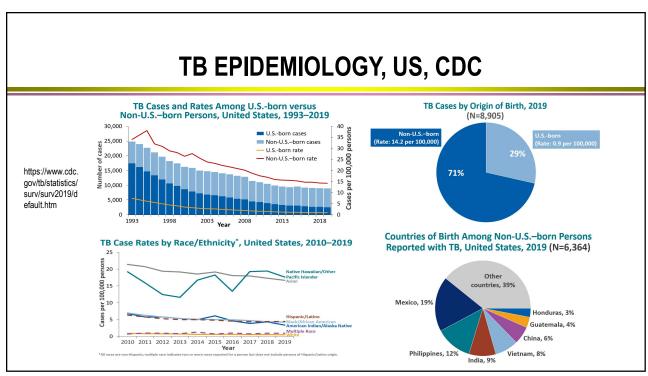
TUBERCULOSIS: THE FACTS

- □ Tuberculosis primarily a disease of the industrial age
- □ Common cause of chronic illness
- □ Well described by term "consumption"
- □ Transmission via airborne route
 - Coughing in source increases transmission frequency
 - Infection control: Mask on source case, detection & therapy LTBI
- Reactivation TB
 - Hemoptysis, 25%
- □ No effective therapy until mid-20th century
- □ Highly effective vaccine still lacking

TUBERCULOSIS TODAY

- □ Falling incidence in the U.S.
- □ Increasingly a disease of immigrants
- □ Worldwide concerns of multi-drug resistance (MDR-TB; resistant to INH and rifampin)
- ☐ Growing threat of extreme-drug resistance (XDR-TB; resistant to INH, rifampin, and ≥2 classes of injectable treatment)
- □ Important cause of death in HIV-infected persons







HIV: THE FACTS

- □ Initial survival of AIDS ~18 weeks
- □ Wasting a common feature of AIDS
- $\hfill \square$ Kaposi's sarcoma considered an AIDS defining diagnosis
- $\hfill \square$ HIV infected persons did in fact face discrimination
- □ Initially no available therapies

HIV: FICTION

- □ Long term survival of persons with HIV infection now common due to multi-drug treatment
- □ PrEP and PEP now available
- □ HIV infected patients protected, in part, from discrimination under the ADA
- $\hfill \square$ Opportunistic infections much less common due to prophylaxis
- □ Kaposi's sarcoma due to HHV-8
 - Incidence has dropped dramatically
 - But still a major public health concern

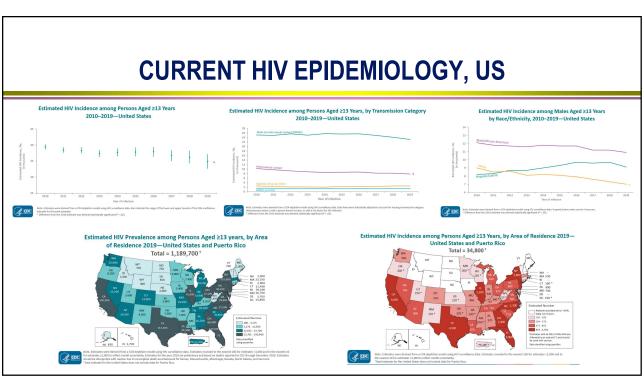




 37.7 million [30.2 million–45.1 million] people globally were living with HIV in 2020.

- 1.5 million [1.0 million–2.0 million] people became newly infected with HIV in 2020.
- 680 000 [480 000–1.0 million] people died from AIDSrelated illnesses in 2020.
- □ 27.5 million [26.5 million–27.7 million] people were accessing antiretroviral therapy in 2020.
- 79.3 million [55.9 million–110 million] people have become infected with HIV since the start of the epidemic.
- 36.3 million [27.2 million–47.8 million] people have died from AIDS-related illnesses since the start of the epidemic.

https://www.who.int/data/gho/data/themes/hiv-aids https://www.unaids.org/en/resources/fact-sheet





IMPACT OF 1918-19 INFLUENZA AND COVID-19 PANDEMICS ON DEATH RATES AND LIFE EXPECTANCY, US

