Right under our noses Emerging & re-emerging Diseases

By Nancy Dewhirst, RDH, BS

Standing up to Emerging Diseases

Drug resistance **Biofilm diseases** Vector-borne diseases Bloodborne diseases Airborne diseases Precautions, prevention & planning What's on the horizon? Outbreak: sudden escalation in cases, local or widespread Epidemic: Widespread, 1 or few areas Pandemic: country-wide or worldwide Affects large % of population Often novel pathogen, no immunity More deaths Social disruption, economic loss Endemic: imbedded, constant or cyclical Safety in perspective "recent" pandemics & epidemics that changed safety standards 1918-1919 "Spanish flu" A(H1N1) 1957-1958 "Asian flu" A(H2N2) 1968-69 "Hong Kong flu" A(H3N2) 1980's HIV / AIDS Bloodborne Pathogen Standard 2002-4 SARS (coronavirus) 2009 "Swine flu" A(H1N1) 2013 Avian flu A(H7N9) 2019 SARS-CoV-2 Respiratory protection standards **Emerging Diseases** Last 20 years: global pandemics Ebola - West Africa,

Zika virus - Americas, Yellow fever - Angola, many are MDR pathogens, COVID Polio - Nigeria: 2016, NY: 2022 TB - Cambodia: 2017, U.S. surge 2023 post COVID

Factors impacting Microbial Pathogen emergence

Increased urban populations Easy transmission

Rapid, unplanned city growth War, refugees Crowding Poor sanitation & healthcare

Global demand for meat

Industrial farming: microbes become pathogens Use of colistin in livestock in China mcr-1 gene Livestock near wild birds largest influenza pandemic in history

Microbial emergence factors

Global warming

Thicker blanket of carbon dioxide allows insects and other vectors to enter new locations

Mosquitos & ticks, bats (vectors, reservoirs)

In a world of microbes, which ones are our enemies?

You are microbial!

Microbiome: "collection of bacteria, fungi & other single-celled organisms" You have 10 X more bacterial cells than human 400 X more microbial genes than human Most microbes are biofilm dwellers & protective if in balance Virome: you have ~10 x more viruses than bacteria Bacteria & viruses - symbiotic

Once Upon A Time, There was the Antibiotic Golden Age......(45 yrs)



Man ID'd, studied & targeted specific pathogens

Broad-spectrum antibiotics

And then...... Pathogens developed resistance Rise of chronic, recalcitrant infections Cystic fibrosis (CF) Pseudomonas aeruginosa Meanwhile.... Researchers found environmental mixed communities of microbes vs. Isolated single organisms

Now know: BF costs billions of dollars / year

energy losses equipment damage product contamination medical infections

A Brief history > 100 antibiotics now used, but almost no new ab's since 1987 Over-use of antibiotics & treating livestock has increased microbial resistance & drug allergies 2016: mcr-1 gene found in humans & pigs (gene creates superbugs) A Brief history Tetracycline, erythromycin, vancomycin = often ineffective All surgeries; implants, transplants, cancer treatment..... Rely on antibiotics WHO ID's drug resistance as major human threat MRSA Multi-Drug Resistant Staph. Aureus Staph = common in flora of skin, nose, throat MRSA colonizes 1/3 of pop. 64% more likely to die than non-colonized

Usually non or mildly infective unless enters bloodstream

Enters through open skin

Pimples, boils, lesions at point of entry MRSA forms biofilms & spreads Infection may lead to pneumonia, severe skin, bone, bloodstream infections, septic arthritis, endocarditis, deep abscesses, toxic shock Drug Resistance: challenges protocol & TX Incidence linked to exposure, susceptibility & over-use of antibiotics Resistance requires more toxic, dangerous drugs Need IV instead of oral meds

Targeted IC had improved MRSA control in hospitals until COVID (crisis lowered control)

Resistant skin infections.... Watch Points of entry & signs of systemic infection MDR Gonorrhea U.S. Cases rising since 2009 Highly resistant strains becoming more prevalent 63% increase last 5 yrs ~ ½ are reported 75% cases = male

MDR Gonorrhea U.S. Cases rising since 2009 Asymptomatic in 80% females & 50% males Gonorrhea - one of 1st pan-resistant diseases (totally uncurable) Prevention: same as HIV Does not respond to cephalosporins Vaccine in development Oral, Ocular Gonorrhea Sore, hoarse, throat Or asymptomatic Eyes = painful, red suppuration (Usually both eyes)

Chlamydia Most common bacterial STD Mostly teen females (immature mucosal columnar cells: vulnerable) 90% asymptomatic Chlamydial Pharyngitis No identifying lesions Fever, flu-like symptoms Wear your eye protection! Screen patients: take temp

Ocular Chlamydial Conjunctivitis Red eye Mild discharge, Foreign body sensation, tearing Blurred vision Swollen eye lid Usually single eye Pre-auricular nodes swollen Usually direct contact Tx: oral doxycycline 100 mg -1 month Syphilis 25% increase – 2015-2019 Many cases multi-drug resistant Primary lesion: chancre Highly infectious spirochete

Emerging resistant diseases Aspergillosis Aspergillus (mold, fungus) – inhaled, usually over time 60% mortality rate Damp buildings, soil, seeds, damp, decaying vegetation Candidiasis (fungus) Candida auris Hospital infection – breathing, feeding tubes, IV lines Emerging resistant diseases Candida auris (yeast, fungus) discovered – 2009 Colonizes skin, room surfaces Resistant to antifungals Causes sepsis 2021: 1st totally resistant case **Shigella (bacteria)** fecal- oral, contact

MDR E-coli (bacteria)

Contact, ingestion

What's the point? Drug resistance: Makes treatment less successful Increases importance of prevention Antibiotic stewardship Creates need for new strategies: Tethered macrocyclic peptides, a new class of antibiotics treat drug-resistant Acinetobacter baumannii Biofilms: the most successful form of life Biofilms: This is how most microbes live

• Structured heterogeneous (mixed) communities (bacteria, fungi, protozoa, nematodes)

Preferred bacterial state (over planktonic)

Homeostasis: stabilize pH, temp., minerals, food, O₂ concentrations

• Aerobic - anaerobic zones

Interdependence & communication

Biofilms

• Protective polymeric matrix:

• 50% - 80% = matrix

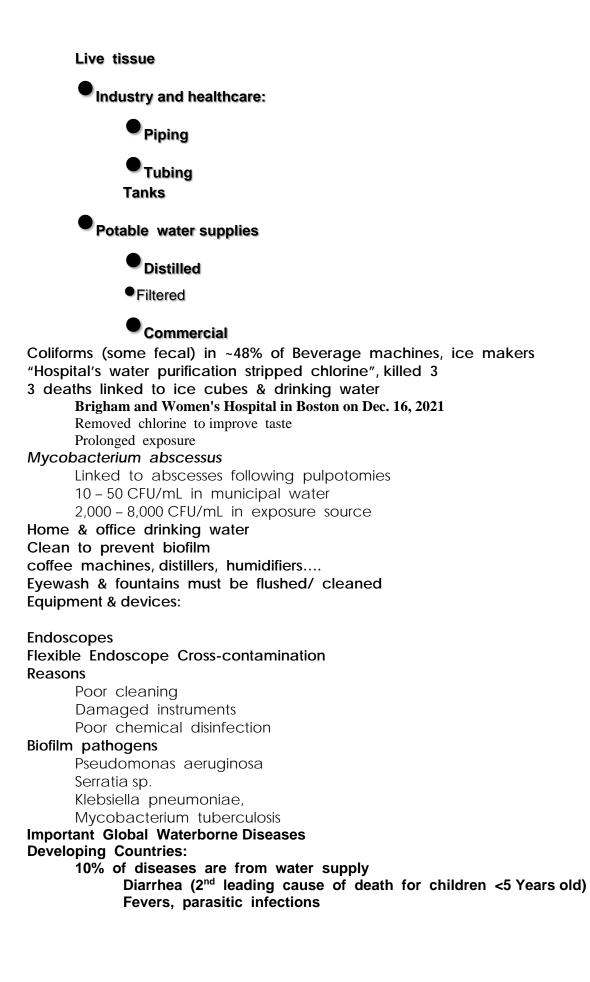
20% = microbial

Bacteria replicate \rightarrow clonal groups

•Some die, leave holes

Pathogens attach to pre-existing biofilms..... Where Do Biofilms Form?

> Widespread in nature lakes streams mud (Require moisture, nutrients, a surface) * Key role in food webs Rhizosphere



Bacteria, viruses, Intestinal worms

No safe drinking water Waterborne Diseases here

• Developed Countries:

- Facing drinking water crisis.
- Decrepit equipment
- Population
- Need for upgraded filtration & treatment
- Extreme weather events

• "Safe water can no longer be taken for granted..." (American Academy of Microbiology)

Waterborne Diseases Developed Countries

•Waterborne pathogens increase significantly each year.

• Pathogens = resistant to treatment chemicals.

- Cryptosporidium
- Giardia
- Cyclospora
- •People live longer, immunocompromised

Requires multi-discipline approach
 Cryptosporidium
 Most common non-viral cause of diarrhea
 Waterborne parasite
 Cryptosporidium ("Crypto")
 Spore phase resists chlorine, desiccation, lives 10 days

Cryptosporidium Causes watery diarrhea Causes 54% of outbreaks (pools & water parks) (CDC) Spread by fecal contamination Don't drink pool water Keep ill swimmers home! Take frequent bathroom breaks and wash! E. coli most common The remaining cases: 75% = Pseudomonas aeruginosa
25% = Legionella
Heat dissipates disinfectants
Physical removal works best but may not be possible
Factoids about biofilms
(key to managing, using & controlling biofilms everywhere)
Commensal behavior within biofilm = protective

• Outer layers absorb antimicrobial agents - outer cells "warn" inner cells to initiate stress response

Stress response: cells change activity & form

•This alters targets of immune systems or chemicals

Persister cells (spore-like) survive = viable but non-culturable: <u>undetectable by</u> tests used

Target: Matrix variations Biofilm & Inflammation wars

Inflammation: standoff between phagocytes & bacteria

Can't penetrate BF

Phagocytes can't engulf biofilm – shoot enzymes at it

Causes general destruction (collateral damage)

Inflammatory response to biofilm infections = heightened, chronic, locally destructive

Biofilm Diseases
Native valve endocarditis =

Biofilm "vegetation" (extremely high concentrations of antibiotics can cure in 6 weeks)

Osteomyelitis

Toxic shock - vaginal biofilms, tampons (staph)
Cystic fibrosis: pseudomonas & mucus
UTI's

Diagnosis = difficult

Cultures only grow when biofilm sheds!

Biofilm Diseases Biofilms in Chronic Wounds > 60% chronic wounds have biofilms Poorly revealed with cultures 2 - 15 bacterial species present

Fusobacterium, bacillus, actinomyces, staph, strep, pseudomonas, candida, 3 types of treponemes Fungi (yeast), no viruses addressed! Biofilms prevent healing Diabetic, venous, pressure ulcers MUST physically remove biofilm to treat Signaling research & challenges Research goal: hijack & control guorum sensing: gain control Challenge: Protect beneficial bacteria and biofilms Keeping signal localized on target Signaling: quorum sensing Bacteria communicate chemically within biofilm (hormone-like signals) Allows biofilm to act collectively as a single force: Aggregate Swarm Disperse Secrete Absorb Replicate Change cell phase (spore, dormant....) Signaling action plans Goal: manipulate signaling to: Prevent formation Slow arowth Trigger detachment (swim away) Challenges: Preventing signaling to beneficial bacteria and biofilms Keeping signal localized on target Bacteria use electrical charges, along nano-wire network within biofilm Maybe: disrupt biofilm communication through power grid **Biofilm Research** Probiotics: designing the "optimal" protective biofilms Probably different for different people Controlling inflammation Will Biofilms Save Earth? "Slime curtains" can isolate & separate toxic areas; protect ground water, streams, rivers Biofilm layer can prevent penetration of acid water drainage into under ground aquifers Engineered biofilms can metabolize toxins, accelerate bio-degradation of wastes Recap: **Biofilm Property Clinical Implications** Behaves as primitive multi-cellular organism Formation is orderly

Cell- to - cell communication required

Target weak links

Target early steps

Find antagonists to intercept, or control signals Biofilm Property Clinical Implications Biofilms resist host immune responses & antimicrobials

Biofilm phenotype = different from planktonic

Biofilms use & respond to electrical signals

No one method found successful Target and remove protective matrix + symbiotic species

Target correct phenotypes

Use electric currents to weaken / disrupt microbes in biofilms, then use antibiotics, antimicrobials

Combine physical, chemical and signaling strategies **DUWL Microbes** Pseudomonas sp. Pasteurella sp. Micrococcus sp. Klebsiella Legionella sp. Mycobacterium sp. Enterococcus sp. Actinomyces Salmonella Strep. ,Staph. **Bacteroides** E. coli Nematodes Protozoa, amoebas Fungi (Candida, Aspergillus sp. **Associated Illnesses** Head, neck, dental infections Septicemia HCA surgical infections Pneumonia, Bronchitis Legionellosis

Abscesses Appendicitis Salmonella poisoning Cryptosporidiosis DUWL - Related Death (2011) Lancet 82-yr old Italian Woman Legionnaires' dis (L. pneumophila) Proven from dentist's waterlines No other exposures 2 Standards for Water Safety Sterile - for surgery, (cutting bone, normally sterile tissue) 0 CFU/mL of heterotrophic water bacteria CDC special update, OSAP, Dental Board law Potable - for non-surgical procedures -500 CFU/mL of heterotrophic water bacteria (meets EPA safe drinking water standards) CDC, OSAP, EPA, Dental Board For Potable Water Your office should: • Flush lines in AM for 2 min./line (handpieces off) Flush lines between patients for 20 sec. Add antimicrobial product to patient treatment water Shock periodically - remove attached biofilm Follow Manufacturer's directions for use (dental unit & DUW product) Monitor water (test) Treat, Shock, and Test ALL waterlines **Biofilms in Buildings** HVAC filters, traps & seams become sources of biofilm Moisture Intrusion Cascade Moisture intrusion \rightarrow Mold growth Bacterial invasion Insects Rodents
amplification, spread Secondary Bacteria, mold, viruses Rot, chemical, microbial off-gassing Eco-system perpetuates itself, further damages structure (holes, deterioration)

More water

Vector-transmitted diseases Zoonotic & vector-borne diseases = increasing with global warning & contact with nature Aedes aegypti mosquito Black-legged tick transmits Lyme disease (considered climate indicator) Hemorrhagic fevers vector-borne Diseases Malaria, Dengue, Zika, Yellow fever, Lyme, West Nile, chikungunya Aedes scapularis now in Florida: yellow fever! Primarily vector transmitted - treat as bloodborne disease Be alert Aedes aegypti mosquitos Aedes aegypti mosquito City dweller, loves humans Can breed in a capful of water Serial biter - rapid spread of pathogens if infected Lyme Disease Ticks carry Borrelia burgdorferi **Headaches** Fever Fatigue Rash Chronic biofilm disease Vaccine in development Lyme Disease Borrelia forms: Cystic form Pleomorphic Granules **Spirochetes** Found in biofilms in joints, brain, Dementia, mental illness Gene editing may work! (CRSIPR/Cas9) Lyme disease vaccine Ticks: population explosion Lyme disease rose 357% in rural areas, 65% in urban (2007 to 2021) Ticks expanding areas - global warming Antibiotic tx works if early Previous vaccine discontinued in 2002: little demand Pfizer's VLA15: 3-dose regimen over 5-9 months, booster 12 mos. Later Targets Borellia burgdorferi Goal: submit to FDA - 2025 polio

1980's - eradicated in U.S.

July/August, 2022: 1 w/ paralysis Tip of the iceberg Don't forget iron lungs Unvaccinated children! Oral-fecal transmission Wild-Type vs. Vaccine-derived Polio Wild virus is not vaccine related Vaccine-derived polio virus: Outside U.S.: oral (Sabin) polio vaccine - live attenuated virus Can become contagious, infect unvaccinated people Low cost, no needles, robust immunity, administered to mouth, portal of entry U.S. Uses injectable (Salk) inactivated virus since 2000 3-4 shots before age 6 = 99% effective vs. paralysis Boosters now recommended to healthcare & wastewater workers ALL unvaccinated people should get vaccinated Respiratory syncytial virus (rsv) Incubation: 4-6 days Symptoms: Runny nose Cough, sneezing, wheezing Fever Low appetite Common under 2 yrs of age Recently in all ages, may be more serious No treatment (relieve symptoms) Vaccine available norovirus Most common cause - acute gastroenteritis in U.S. Symptoms: extreme vomiting & diarrhea Most common Nov. to April (but year around) Ingestion: food, water, hand-to-mouth (restaurants), recreational & drinking water Infective dose: <100 virions. III people shed billions even >2 weeks after symptoms resolve No vaccine, hand sanitizers not effective Zoonotic transmission to humans Coronaviruses - bats Hantavirus – rodents Influenza - birds, pigs Ebola – primates, bats Insect-borne diseases Parasitic infections Antigenic shift: two viruses (1 from human) share DNA in a common host (influenza a) Cutaneous leishmaniasis Tropical disease Endemic in Middle East, Asia, Africa, Latin America

Global warming: now endemic in U.S. Caused by parasites spread by sand fly bites

Cutaneous leishmaniasis Facial scarring for life Depression, stigma

Aggressive invasive group A streptococcus 2023-4 surge

Strep commonly infects throats, skin

Hyperimmune responses: sepsis, scarlet fever

Invasive gp. A: spreads to bloodstream, meninges (meningitis), organs, soft tissue (necrotizing fasciitis)

"Flesh-eating Strep"

Global increase in cases: 50% - 200% more cases

3/4 cases - hospitalized, ~10% fatal

Aggressive invasive group A streptococcus 2023-4 surge

Severe sore throat & fever, no cough or cold symptoms

Fine, sandpaper-like skin rash, extreme lethargy

Often with flu, chickenpox, COVID-19

Antibiotics needed early

Mpx - zoonotic pathogen

Endemic in West Africa – prior to now: direct exposure to bush meat & rodents, limited person – person transmission

2017-2020 increased cases in Nigeria linked to:

Deforestation, exposure to forest sources

Waning smallpox immunity (1970's vaccinations)

2022: world transmissions - novel event

Spread domestically, (intimate contact, children, pets)

U.S. Had largest MPX outbreak in world (>28,000 cases)

Мрх

Mpx virus

Signs / symptoms: fever, chills, headache, muscle ache, lymphadenopathy, then rash, vesicles, pustules, umbilication, crusts - on face, hands, trunk, genitals Some recent cases start with rash, seen in gay community

Spread by body fluids, close & intimate contact, close-up respiratory droplet exposure (throat lesions shedding), surfaces, materials (bedding)

Port of entry: broken skin, mucosal /ocular absorption

Most infectious phase = rash, lesions: confused with STD's: herpes, syphilis, varicella zoster

Мрх

infectious until lesions totally resolved: new skin formed

Smallpox vaccine

Attenuated virus

Only effective prior to symptoms

Multiple puncture technique with bifurcated needle

Adverse effects common:

Papule at vaccination site (2-5 days) Inadvertent inoculation of other sites **Requires boosters** What about vaccines? How Do We Combat Fear & dis-information? With science & logic Vaccine basics: All vaccines: ~5-10% of vaccinated may not respond (or weakly) Vaccines assist immunity, Build antibodies ~ 2 weeks Host's immune system determines the strength of both recovered (convalescent) & vaccine immunity Immunocompromised likely to have less & shorter immunity science & logic: Vaccine basics Vaccines prevent infection Polio Measles Smallpox Vaccines may prevent infection, or REDUCE SEVERITY Influenza, COVID Pathogens mutate Vaccines do not make PPE unnecessary Vaccines are specific to one pathogen What about the others? **Bloodborne Diseases** (blood & fluids = infectious) Examples: HIV, hepatitis Hepatitis B CDC 2023 Updates Screen all >18 years at least once - triple panel test HBsAg = chronic or acute infection (or recent vaccine, temporarily) Antibody to HBsAg (Anti-HBs) – indicates infection recovery, indicates immunity in never infected vaccinated Total antibody to core antigen (anti-HBc) indicates HBV infection, lasts for life Chronic infection: total anti-HBc & HBsAg (+) HBV DNA measures viral load HBeAg indicates viral replication, high infectivity Hepatitis B CDC 2023 Updates Screen all pregnant, each pregnancy for HBsAg Regardless of history of tests or vaccine **Risk-based testing for:**

Incarcerated Multiple sex partners HCV (+) Test anyone who asks for test

5

https://www.cdc.gov/mmwr/volumes/72/rr/rr7201a1.htm?s_cid=rr7201a1_w

Ebola Virus **5 Enveloped Viruses** 4 infect humans **Ebola Virus Transmission** Direct contact with all body fluids / substances of a symptomatic person or animal (bats, bush meat) Blood, urine, feces, vomit, sweat **Ebola Transmission** Exposure to contaminated objects (sharps: needles = highest risk...) Sexually transmitted, > 1 year after male recovers PPE to Treat Suspected / known Ebola patients Full body suit, no skin exposed **Double gloves Fit-tested** respirator Training to safely don, remove, use PPE Trained helper to don & remove PPE Ebola Survivors' Ailments Neurological symptoms (~75% of pts.) Memory loss, cognitive disorders Headaches, Parkinson's-like symptoms Extreme fatigue, anxiety, depression, sleep disorders Eyes (~60% of pts): Cataracts, blurred vision, redness, pain, light sensitivity, detached retina, blindness, light flashes Muscles: pain, weakness Joints & cartilage: pain Ears: ringing (tinnitus), deafness / hearing loss Ebola (like HIV) hides in reservoirs & recurs in episodes Virus hides in reservoirs with "immune privilege" Eves Testes (> 1 year after recovery, even with (-) blood test) Joints, joint cartilage Brain Uterus? Immune (macrophage) response to Ebola virus may set off cytokine storm in brain, joints (like HIV) fatigue, pain.

Ebola vaccine: Ervebo STRIVE = rVSV-ZEBOV (recombinant Vesicular Stomatitis Virus Zaire ebolavirus vaccine) approved in 2019 Protects against Zaire ebolavirus Hindered by fear!!! Vaccine cannot cause Ebola because it does not contain the whole Ebola virus Single-dose injection, and is a live, attenuated vaccine Education needed > $\frac{1}{2}$ of polled Africans still have mis-conceptions: "mosquitos or ambient air spread Ebola" Majority improved handwashing & touching suspected Ebola victims Burning clothes, belongings of patients Ebola = Category A infectious substance No products specifically list Ebola Use high potency registered disinfectant with label claim vs. Non-enveloped virus (norovirus, toravirus, adenovirus, poliovirus) Ebola = enveloped virus Margin of safety: will inactivate both classes of viruses Microbial resistance to killing Prions **Bacterial endospores Fungal spores** Mycobacteria - Mycobacterium tuberculosis Nonlipid or small viruses (Non enveloped) - Polio virus, enteroviruses Fungi - Trichophyton spp. Vegetative bacteria - Pseudomonas aeruginosa, Staphylococcus aureus Lipid (enveloped) or medium-sized viruses - Herpes simplex virus, hepatitis A, B & C virus, HIV, Ebola, SARS-CoV-2 (CDC) **Follow Label Directions Clean before disinfecting Proteins neutralize disinfectants** Infectious diseases Bloodborne diseases are critical, but.... 80% of common infections (colds, flu, diarrhea) - spread by contact, air, water, food, fomites Stay informed: ADA.org, CDC.gov, OSAP.org, AGD.org, CDA.org

Aerosol Transmissible Diseases

Pathogens capable of surviving air suspension:
Desiccation
Travel on dust particles, air currents
Particles < 5*u* large enough for viral load, small enough to travel > 20'
Absorbed through conjunctivae, mucosal tissue of nose, respiratory tract
6' distancing is not enough

Covid concerns

New variants – evade immunity

Current surge (detected in wastewater)

Long COVID???? (1 month after illness) ~10% of U.S. cases

Fatigue, respiratory, cardiac, neuropsychiatric and GI dysfunctions.... Immunocompromised, diabetes, heart, lung, kidney diseases... Autoantibodies

Covid concerns

SARS-CoV-2 linked to newly diagnosed diabetes & heart damage & attacks, arrhythmias, strokes, clots

 \geq 30 days after infection All ages! (Not just <18)

Post-COVID (even mild); screen for:

Frequent urination, increased thirst & hunger, weight loss, fatigue, stomach pain, nausea, vomiting

Arrythmias, heart attack & stroke symptoms

Ask patients & be self-aware

Criteria for determining risk

is it safe????

Disease activity locally

Specific pathogen features (mode of transmission, transmissibility, severity) Mitigation strategies in place

Eliminate/reduce contact & exposure

Tele-dentistry, distancing, barriers

Engineered safety devices / technology

Suction, HVAC, Air filtration & changes

Rules, protocol, management (screening, source control...)

<mark>PPE</mark>

Vaccination status + immune profile Aerosol generating procedures Airborne Transmission of SARS-CoV-2

Influenza Highly infectious respiratory disease Epidemics since 1510 Mild to severe, usually targets very young, elderly, weak All flu viruses originate in birds most stay there

SARS-CoV-2 has changed dental safety standards Consider everyone infectious for ALL types of diseases, including aerosoltransmitted diseases Plan for safer buildings, more air management Upgrade traditional PPE Exposure response Apply today's lessons to your healthy future! Understand each type of disease Use appropriate precautions Right under our noses Emerging & re-emerging Diseases

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