

Respiratory Viruses, How They are Transmitted, and the Potential Role of Surfaces

Author:

Peter Teska, MBA; James Gauthier, MLT, CIC; David Buckley, PhD

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Abstract

While respiratory viruses are a global health concern, significant gaps exist in our knowledge of how they are transmitted, including the portion of transmission that occurs via contact, droplet, and airborne routes for a given respiratory virus. This paper reviews the current evidence of transmission for a range of respiratory viruses. The studies discussed indicate that contact transmission is an opportunistic mode of transmission for respiratory viruses and that hand hygiene and surface disinfection are likely to play a role in reducing transmission risk.

Background/Introduction

Respiratory viruses are a global public health concern. This is no more evident than during the COVID-19 pandemic, which has created significant morbidity and mortality, and will likely be remembered as the greatest public health crisis in the last 100 years.

Despite >200,000 scientific papers about SARS-CoV-2 being published in 2020 alone (Else, 2020), fundamental gaps exist in our knowledge about the virus and the disease it causes, often leading to disagreement between medical experts, public health experts, and scientists, impacting consensus recommendations to help contain the pandemic. Gaps in our knowledge of SARS-CoV-2 have prompted a review of data about other respiratory viruses to determine whether studies done on other respiratory viruses, such as influenza and measles, might better inform our understanding of SARS-CoV-2.

One of the most important epidemiological questions concerning respiratory viruses is understanding how they are transmitted between people. As is discussed below, it is clear from the literature that there are three transmission routes of concern for respiratory viruses: droplet, airborne, and contact (including fomites/surfaces) (CDC, 2018) (CDC, 2019) (CDC, 2020) (CDC, 2021). However, there is little agreement of the portion of transmission that can be attributed to any one mode of transmission for a given respiratory virus and there is some evidence that the portion of transmission by mode is likely different between respiratory viruses (Lei, 2017) (Otter, 2016).

Fundamental to stopping the spread of any pathogenic microorganism is understanding the epidemiology, which includes understanding the routes by which the pathogen is spread between people, commonly referred to as the "chain of infection" (see below). By understanding how the pathogen is spread, preventative

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measures can be proposed that would interrupt transmission. By understanding the portion of transmission occurring via a given mode, resources can be targeted towards interrupting the modes likely to cause the majority of transmission.

Key questions that need to be answered in order to provide more clarity on respiratory virus transmission are listed below. For some of the questions, there is already some evidence, but typically the evidence is limited in scope to a small number of trials, which may not be representative across various populations and different settings.

- Finer resolution regarding percent of people that are asymptomatic throughout infection
- Stronger evidence regarding distance between people that is necessary to acceptably reduce risk and how other interventions (i.e. barriers or ventilation) being present affect this risk
- Parsing out transmission via air: airborne versus droplet
- Protective effect provided by nonrated (i.e. homemade) masks versus rated surgical masks in preventing infections
- Protective effect provided by eye protection
- How much of transmission is related to fomites (surface) and what factors affect this portion of transmission

The US Centers for Disease Control and Prevention (CDC) (2021) recently published a science brief on the risk of fomite transmission for SARS-CoV-2 to build on prior CDC guidance regarding transmission of SARS-CoV-2. This document rightly indicates that exposure to respiratory droplets in close contact are believed to be the dominant mode of transmission, but considers surface mediated transmission a low risk. Because of the difficulty in quantifying the portion of transmission that occurs via any one route and because of the important role of surfaces in the transmission of a wide range of other pathogens, including other respiratory viruses, this position has been controversial.

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The aim of this paper is to address the risk of environmental transmission of respiratory viruses and review the available evidence that supports various modes of transmission. The studies discussed below indicate that contact transmission is an opportunistic mode of transmission for respiratory viruses and that hand hygiene and surface disinfection are likely to play a role in reducing this risk when it does occur.

The "Chain of Infection" Model:

Human pathogens are those microorganisms that cause disease in people. For pathogens to survive, they must move from one host to another or to a reservoir over time. The "chain of infection" model, developed by the Centers for Disease Control and Prevention (CDC), describes the key factors in transmission of pathogens that can result in infection (CDC, 2012). The factors include:

- The **infectious agent** (i.e. the pathogen causing disease)
- The reservoir for the pathogen, such as an infected person or animal carrying the pathogen
- A **portal of exit** from the reservoir for the pathogen to start the transmission process
- The **mode of transmission** to move from the reservoir to a susceptible person
- The **susceptible person** that is at risk of being infected, and
- A **portal of entry**, where it gains entry into a susceptible person and thus initiates infection

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Figure 1. Chain of Infection / Transmission Model (Disease Detectives, n.d.)





Any discussion on how infectious diseases are transmitted would include a focus on the key roles of the **reservoir**, the **susceptible person**, and the **method of transmission** as part of the chain of infection, as these are the primary targets of intervention programs designed to prevent infection (CDC, 2016).

For specifically. respiratory viruses eliminating the pathogen reservoir for the virus can prevent transmission by reducing the amount of pathogen available to cause infection. Reducing a person's susceptibility, such as by wearing a mask or receiving vaccination, reduces a person's risk of being infected by a respiratory virus. Preventing transmission of the pathogen, such as by mask, social distancing. wearing а performing hand hygiene, or disinfecting surfaces all may interrupt the chain of infection and thus prevent disease. Of the three, eliminating the pathogen reservoirs the method of transmission for and respiratory pathogens are the focus of discussion for this paper.

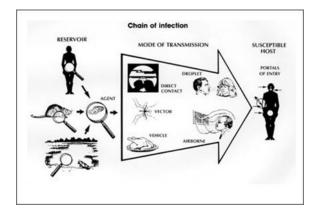
Modes of Transmission

The standard CDC reference document on pathogen transmission is the Siegel (2019) guidelines originally published by CDC in 2007, which lists these modes of transmission: <u>Contact</u> transmission: divided between **direct** and **indirect** contact, where microorganisms are either transmitted from one infected person

- to another without a contaminated intermediate object or person (i.e. direct contact) or by a process involving an intermediate object or person (i.e. indirect contact). Vectorborne transmission (i.e. contaminated food or water or through an animal or insect carrier/reservoir) is another type of contact transmission and some other references refer to this as a separate type of transmission.
- <u>Droplet transmission</u>: involves respiratory droplets carrying infectious pathogens. In this context a distance of 3-6 feet would be high risk with droplets >5 microns as likely to transmit infection.
- Airborne transmission: involves respiratorv droplets carrving infectious pathogens over a longer distance or for a prolonged period of time. The guidance does not state specific distances or droplet sizes as requirements for determining airborne transmission, but generally it is assumed that smaller droplets (i.e. < 5 microns) and longer distances can be involved. Smaller droplets can stav airborne for longer periods of time and droplets smaller than 5 microns can be inhaled deeper into the respiratory tract, both of which may make transmission resulting in infection more likely.
- <u>Transmission from the Environment</u>: includes microorganisms commonly found in soil and water. In this type of transmission, other people and intermediate vectors are not involved in the transmission and the pathogen is not transiently contaminating the reservoir but rather is found in the reservoir on an ongoing basis.

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Figure 2. Modes of Transmission for Infectious Diseases (CDC, 2012)



We note that transmission of a respiratory virus can simultaneously occur via multiple modes of transmission. For example, an exposure event can involve inhalation of virus and hand contamination leading to selfinoculation. If studies were done to determine the amount of virus acquired by each pathway, a determination could be made about the significance of each mode of transmission in a specific exposure event.

In the current COVID-19 pandemic, much of the discussion on airborne versus droplet transmission for SARS-CoV-2 focuses on droplet size and which droplets are more likely to carry active virus. Sidestepping much of this debate, we briefly note that there is accumulating evidence that smaller droplets (<5 microns) are more likely to carry infectious virus than larger droplets and that these smaller droplets can travel further than 2 meters (6 feet) in certain circumstances as summarized by Tang (2021), but most studies on transmission show prolonged close contact was a significant factor in transmission, making droplet transmission probable overall а more mode of transmission than airborne transmission (Meyerowitz, 2021).

Chen (2020) provides a detailed discussion of the various sub-routes for close contact transmission and the variables involved.

Their modelling suggests a decrease in risk with increased distance and that risk is heavily impacted by how the virus is disseminated (talking versus coughing, etc.). They also found there was limited risk for those who were distanced by >1.5 meters, regardless of the droplet size or virus dissemination activity, suggesting droplet transmission is more common than airborne transmission. The interconnectedness of modes of transmission is an important concept to note and factors into the surface discussion on transmission. Brankston (2007) states that few respiratory viruses are thought to be exclusively transmitted via a single mode of transmission and challenges in controlling confounding experimental variables make desians complicated.

Respiratory Infections and Mode of Transmission

Respiratory infections, or infections involving the upper or lower respiratory tract, are typically caused by viruses such as rhinovirus, respiratory syncytial virus (RSV), influenza. parainfluenza, human metapneumovirus. measles. mumps. adenovirus, and coronaviruses (LaRocque, 2019), but can also be caused by bacteria, such as Bordetella pertussis (which causes Whooping Cough), or fungi, such as Aspergillus. For this discussion, the focus is on viruses causing respiratory infections.

In a common upper respiratory infection, where the pathogen infects the respiratory tract tissues above the lungs, common activities such as breathing, coughing, sneezing, talking, singing, and shouting can all expel respiratory droplets into the environment which contain the virus (Bischoff, 2013). In a respiratory infection, the virus replicates in the respiratory tract. Both infectious and non-infectious particles are expelled via saliva, mucous, and other respiratory secretions in droplets ranging in

size from <1 micron to 500 microns (Kutter, 2018) (Otter, 2016). How this process occurs is generally understood, but many of the mechanistic details critical for preventing infection are poorly understood, which is where the controversy occurs.

How the body expels a respiratory virus may cause significant variations in the viral load. Tang (2014) discussed the difference in respiratory secretions for people infected with influenza and the impact it may have on sampling, stating that virus mixed with saliva and mucus in the oral cavity is not of uniform viscosity and lower viral load fluids are likely to contain more saliva and thus be of lower viscosity. Saliva containing fluids may be preferentially expelled from the body which could explain lower viral levels in expelled breath in some studies (Tang, 2014). Tang (2014) also discussed that large droplets are mostly generated from the front of the mouth and respiratory viruses, such as influenza, which are less commonly found in saliva, may be less likely to be detected if a person is expelling primarily larger droplets. Saliva can have antiviral properties and expelling virus in saliva may also reduce the viral load detected from large droplets (Tang, 2014). Studies that review expelled breath or light coughing may then give different levels of virus than studies that capture sneezing or heavy coughing even if study participants had similar viral loads.

Boone (2005) (2007) points out that respiratory viruses can be expelled from the body via sneezing and coughing at velocities of 20-45 m/sec (45-100 mph) and >3 m of distance with concentrations of 10⁷ virions/mL. Sneezing can similarly cause rapid dissemination of respiratory droplets at distances of more than 6 meters.

The concept of viral load is important because modelling studies commonly use studies of viral load to calculate risk of transmission. If the viral load is lowered by TIPS www.IC.tips Join. Contribute. Make A Difference.

the activity used to generate the contamination in the study, this may introduce a bias, decreased the modelled risk and most modelling studies tested for sensitivity identify the viral load as strongly influencing the modelling result.

On the CDC website, there are webpages for many common respiratory diseases, which includes a wide range of information about the virus including the identified routes of transmission. Using this CDC data we can assemble the table below.

Table 1. Common respiratory viruses androutes of transmission

Common Respiratory Viruses	Contact	Droplet	Airborne	References
Adenovirus	Yes	Yes	Not indicated	(CDC, 2019)
Coronaviruses - non-SARS	Yes	Yes	Not indicated	(CDC, 2020)
Coronavirus – SARS related	Yes***	Yes	Yes**	(CDC, 2020)
Human metapneumovirus	Yes	Yes	Not indicated	(CDC, 2019) (Kutter, 2018)
Influenza	Yes	Yes	Not indicated	(CDC, 2018)
Measles (Rubeola)	Yes	Yes	Yes	(CDC, 2020) (Kutter, 2018)
Mumps	Yes	Yes	Not indicated	(CDC, 2021)
Parainfluenza	Yes	Yes	Not indicated	(CDC, 2019) (Kutter, 2018)
Respiratory Syncytial Virus	Yes	Yes	Not indicated	(CDC, 2020) (Kutter, 2018)
Rhinovirus	Yes	Yes	Not indicated	(CDC, 2020)
Rubella (German Measles)	Yes	Yes	Not indicated	(CDC, 2020)

(** = in certain circumstances, ***= infection by this route is listed of low probability for SARS-CoV-2)

In developing these recommendations from the CDC, randomized controlled trials (RCTs) are not cited to determine the relative portion of infection caused by each mode of transmission. Consequently, it is challenging to predict the distribution of infections by mode for any respiratory virus. Quantitative Microbial Risk Assessments (QMRA) often are used to model infection risk, but are inherently limited by the assumptions and underlying data in the model. Ideally these models are a first step in predicting the probable outcomes and RCTs with clinical endpoints would then be run to validate the modelling, but this is rarely done.

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For a respiratory infection to occur (Killingley, 2013):

- Replication-competent virus must be present and survive in a reservoir, and
- Must be transmitted from a reservoir to susceptible permissible host cells,
- In a quantity capable of causing infection (i.e. an infectious dose)

If any of these actions do not occur, then infection is prevented. While SARS-CoV-2 is a novel virus, and thus our knowledge of it is more limited, there are examples in the literature from studying other respiratory viruses which are relevant for this discussion.

SARS-CoV-2 and Modes of Transmission

The World Health Organization (WHO) (2020) states that SARS-CoV-2 is spread by people in close contact (i.e. <1 meter apart) when aerosols or droplets come directly into contact with the eyes, nose, or mouth and that poor ventilation can increase risk. The WHO (2020) also states that people can become infected by touching surfaces that have been contaminated with SARS-CoV-2 and then touching their eyes, nose, or mouth.

CDC (2020) states transmission is thought to spread through close contact (<6 feet, or <2 meters) through respiratory droplets, but that the virus can also be spread in certain situations by airborne transmission, where people further than 6 feet (2 meters) away or for up to several hours after the infected person has left the space. The CDC further states that SARS-CoV-2 is spread less commonly through contact with contaminated surfaces (CDC, 2020), as was summarized in the recent CDC science brief (2021).

Roy (2004) proposed classifying communicable respiratory pathogen transmission as:

- Obligate when transmission **only** occurs via a primary route.
- Preferential when transmission occurs primarily through a dominant route, but other routes are possible, especially under different sets of conditions.
- Opportunistic when for **certain routes**, transmission only occurs under certain sets of conditions.

This language proposed by Roy is helpful for this discussion as the CDC and WHO statements on transmission suggest a primary or preferential mode of transmission (droplet) and secondary or opportunistic modes of transmission (airborne and direct/indirect contact).

The challenge for WHO and CDC was noted by Tang (2021) who commented that there is little direct evidence of transmission of SARS-CoV-2 via any specific mechanism or mode, largely due to the expense and difficulty in performing experiments that would prove this conclusively. Of the SARS-CoV-2 published studies on transmission, they contain lower quality evidence that lack sufficient controls, suffering from confounding, or rely on modelling, which comes with its own limitations.

The remainder of this paper reviews available evidence on respiratory virus transmission, starting with SARS-CoV-2, and compares that to evidence for other respiratory viruses to demonstrate the potential for opportunistic contact transmission and whether there are conditions under which this mode of transmission is more likely.

SARS-CoV-2 and Environmental Survival and Transmission Considerations

As indicated above, the first consideration is whether the virus is likely to remain active for prolonged period of time in the а environment so that opportunities for transmission occur. Testing of the ability of SARS-CoV-2 to remain active on surfaces was performed early in the pandemic by Van Doremalen (2020), who tested SARS-CoV-2 for viral viability in an artificially created aerosol (with similar levels to a person with a respiratory tract infection) and when inoculated on a range of surfaces, including copper, cardboard, stainless steel, and plastic. The authors found that the aerosol remained active for up to 3 hours (half-life of 1.1-1.2 hours) and on plastic surfaces for up to 3 days (half-life 6.8 hours) (van Doremalen, 2020). As this testing involved determining if infectious virus was present, this strongly suggests a transmission risk associated with surfaces for up to several days with the highest risk immediately after the surface is contaminated and a declining risk over time.

Meyerowitz (2021) summarized the available evidence on SARS-CoV-2 and found a wide range of studies detecting SARS-CoV-2 RNA on environmental surfaces. While the level of viral RNA detected on surfaces in studies is typically orders of magnitude lower on surfaces than in nasopharyngeal swabs (Meyerowitz, 2021), this is to be expected since the time between surface contamination, time of sampling, and efficiency are uncontrolled sampling variables in these studies. Presumably the highest risk of transmission via surfaces is shortly after surface contamination occurs, with the risk steadily declining over several days, as indicated by van Doremalen's study (2020).

Meyerowitz (2021) also hypothesized that the dominant (i.e. preferential) mode of

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transmission of SARS-CoV-2 is to be respiratory droplet with proximity to the source and duration of exposure being critical factors. This suggests airborne transmission is more opportunistic in situations where there is prolonged contact, and poor ventilation. crowded areas, Investigations cited in this review article reporting the potential for fomite transmission confounded the are by evidence being circumstantial or the presence of other confounding variables (Meyerowitz, 2021). Hand contamination is believed to occur primarily through contact with contaminated surfaces and studies have shown poor hand hygiene to increase SARS-CoV-2 infection and risk increased disinfection of surfaces to decrease risk, but these studies are confounded by multiple additional interventions being present (Meyerowitz, 2021).

Additionally, studies of the overdispersion constant (k) show that an estimated 80% of secondary transmission is caused by <10% of infected people (Meyerowitz, 2021), suggesting the potential for transmission dynamics of so called "super spreaders" to drive the bulk of transmission. This data is important because it suggests typical conditions are not those that favor significant transmission, but rather the presence of a super spreader changes the transmission dynamics in poorly understood ways. Consequently, studies of typical situations be less informative may on actual transmission risk if the presence of a super spreader cannot be included.

Studies Involving SARS-CoV-2 and Transmission

In considering the level of virus that may be present on environmental surfaces, it is relevant to understand at what level the virus may exit the body. Several studies investigated the level of SARS-CoV-2 present in common respiratory secretions.

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Wyllie (2020) reported virus levels in saliva of 5.09 -6.07 \log_{10} copies per mL and 4.93 \log_{10} copies per mL in nasopharyngeal swabs. To (2020) reported 5.2 \log_{10} per mL for oropharyngeal swabs. Both studies demonstrate significant viral levels of approximately 10⁵ to 10⁶ genomic copies per mL of secretion. These levels of virus are comparable to other respiratory viruses as influenza is typically detected at levels of 10⁵ to 10⁷ copies per mL (Otter, 2016).

A recent study by Yang (2021) at the University of Colorado Boulder of students living on campus in dormitories had saliva samples taken weekly during the Aug-Dec 2020 school semester, which resulted in 72,500 samples being tested. While the distribution of virus detected in infected students formed a Bell-shaped curve centered on 7.3x10⁵ (i.e. 730,000) virions/mL, the student with the highest viral load had 6.1x10¹² (i.e. 6.1 trillion) virions/mL and the student with the lowest viral load had 8 virions/mL. demonstrating an extremely wide range of viral load. The students with the highest viral load had 8.4 million times more virus than the average infected student. When Yang (2021) converted all the samples to a number of virions in the sample. Yang (2021) determined that approximately 2% of infected students accounted for 90% of all virus detected at a given time. This suggests that these high viral load students are much more likely to cause transmission than the average infected student with a much lower viral load and thus potentially are super spreaders (Yang, 2021). Since a person needs roughly 100,000 virions/mL to have any infectious SARS-CoV-2 virus (La Scola, 2020), and thus pose an infection risk to others, this suggests at least half of those infected with SARS-CoV-2 are at no risk of onward transmission any time during their infection and attention should be focused on only those infected people with high levels of virus.

As mentioned previously, one of the unresolved questions around SARS-CoV-2 is whether the virus is transmitted via an airborne route. While there are many advocates for airborne transmission being a dominant route. Meyerowitz (2021)discussed the result of a 40 study metaanalvsis that focused on household transmission which found a secondary attack rate of 18.8%. If SARS-CoV-2 were preferentially an airborne transmission, secondary attack rates would be expected to be >80%, as would typically occur for measles and chicken pox (CDC 2015), suggesting that airborne transmission is opportunistic for SARS-CoV-2.

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A study looking at this question in an indirect method was a study by Zeng (2020), who reported that for a group of people who contracted COVID-19 in a city in China, the percent of people who wore eyeglasses on a daily basis (5.8%) was substantially lower than in the general public (31,5%). suaaestina evealass wearing had a protective effect on SARS-CoV-2 infection. If SARS-CoV-2 was preferentially transmitted as an airborne infection, eyeglass wearing would be expected to have little effect since it provides some protection from ballistic droplets, but not smaller droplets floating in the air, where eyeglass wearing would be less likely to provide some protective effect.

These two studies do not prove airborne transmission is opportunistic, but they provide some evidence supporting this position. Studies investigating the role of surface transmission similarly provide evidence that surface mediated transmission occurs, but it similarly is opportunistic and not a preferred route of transmission.

Wang (2020) performed a retrospective cohort study of preventing secondary attack in a household with a sick family member. As >70% of cases from the initial outbreak in China occurred in families, preventing

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secondary attack was of interest. In the multivariable logistic regression model, transmission was significantly reduced by frequent surface disinfection with chlorine or ethanol and mask wearing by all family members (Wang, 2020). Interestingly, hand hygiene alone did not have a significant impact, but when paired with mask usage the impact was significant (Wang, 2000).

Harvey (2021) performed environmental sampling on surfaces in Somerville, Massachusetts and found 8.3% of surfaces were positive for SARS-CoV-2 RNA with levels ranging from 2.5 to 102 genomic copies per cm². The authors note the potential confounding of routine cleaning and disinfection, hand hygiene, and use of masks, which may impact the levels of virus detected. Based on the sampling, they modelled the risk of contact transmission and infection from a single hand to surface contact as 1 in 100,000 to 1 in 2,500 and noted this is less than the modelled risk associated with influenza on surfaces at 1.25 in 10,000 (Harvey, 2021). The authors note that their model is highly sensitive to the level of virus detected on the surface.

Xie (2020) reported on transmission of SARS-CoV-2 in Guangzhou China, tracing contacts of a cohort of infected people and their movements. In one case due to environmental testing and video of movements and actions, they were able to show that the likely transmission occurred via a contaminated elevator button when the infected person blew their nose into their hands and then touched the elevator button 2020). video evidence (Xie. Having supporting transmission is rarely available to researchers, leading to challenges in performing similar studies.

Moore (2021) reported on detecting SARS-CoV-2 RNA in 8 hospitals in England and found that 8.9% of environmental samples were positive for SARS-CoV-2 RNA, but the

contamination was not widespread with only 10 bed sites accounted for 63% of all environmental sites positive for the virus RNA, suggesting the daily cleaning protocols were largely effective at removing the virus for most bed spaces. None of the environmental samples contained replication competent virus, but the timing between surface contamination and sampling was not explored (Moore, 2021). The highest level of virus detected was 1.6 x 10³ genomic copies per swab on a pulse oximeter, suggesting some patient care equipment may be more heavily contaminated with the virus, especially if a COVID-19 patient contacts the equipment with their hands (Moore, 2021).

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Studies Involving Other Respiratory Viruses and Transmission

One of the points driving this discussion is that while SARS-CoV-2 is novel and thus not well understood, being a respiratory virus helps inform our understanding of how SARS-CoV-2 likely behaves and how transmission is likely to occur. This section reviews select studies of other respiratory viruses.

<u>Influenza</u>: One of the most commonly studied respiratory viruses is influenza, having been studied over a number of decades. Despite this depth of investigation, there remains a lack of consensus on the portion of infections occurring via any specific mode. Five studies are discussed below.

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Table 2: Select Studies InvestigatingEnvironmental Survival of InfluenzaVirus

Reference	Study Location	Inoculation Level or Sample Population	Virus Survival or Detection Comments	Implications or Other Comments
Thompson 2017	Laboratory	10 uL of 10 ⁶ – 10 ⁶ pfu/mL	99% reduction in 17.7 hrs for cotton 174.9 hrs for stainless steel	Rapid inactivation in first hour Viable virus recovered for up to 2 weeks. Surface material had strong impact on virus survival.
Oxford 2014	Household surfaces in lab setting	500 uL of 5.5 log10	8 hours on cloth 48 hours on wood	Recovery on cloth challenging and may have influenced survival time.
Boone 2005	Day care facility and homes	Environment sampled	Day care: 23% of surfaces positive in fall and 53% positive in spring Homes: 59% of surfaces positive in spring, 0% positive in summer	No influenza detected in summer when flu rates are low. Influenza only detected in homes with infected children. Virus detected most frequently on phones and computers.
Bischoff 2013	Acute care facility	Patients with influenza-like illness air samples	65% of patients infected with influenza and 43% of these patients had positive air samples. Up to 89% of influenza particles were <4.7 microns in size.	Patients with influenza had higher viral loads when shedding virus into the environment. 19% of patients shed influenza into the air at levels 32 times higher than other patients. Viral load decreased rapidly from 0.305 m to 1.829 meters from the patient's head.
Asadi 2020	Tissues in lab setting	3.6x10 ⁵ PFU per fomite	Crumpling, folding or rubbing tissue released 0.3 – 10 micron particles with active virus	This demonstrates a potential to re-aerosolize a respiratory virus, such as influenza.

<u>Respiratory Virus Studies</u>: A number of studies have been done on respiratory viruses as a group and six of these studies are summarized below.

Table 3: Select Studies InvestigatingEnvironmental Survival of RespiratoryViruses

First Author Year	Study Location	Sample Population or Study Environment	Virus Survival or Transmission Comments	Implications or Other Comments
Zulli 2021	Elementary School	Student desks	Rhinovirus on 8.9% of desks, adenovirus on 6.7% of desks, coronavirus on 3.1% of desks. Influenza on 0% of desks	Cleaning reduced the level of virus on desks, but returned to precleaning levels in 2-5 days. Sitting at 5 desks per day >60% chance of contacting respiratory viruses.
Phan 2020	Acute care patient rooms	Patients with respiratory infections	Virus causing infection found on 43% of surfaces 22% of air samples. 67% of computer keyboards and 52% of bedrails	Positive correlation between level of virus and number of hand contact for surfaces commonly touched by healthcare workers (HCW) including IV pole, telephone, and bed control.
Otter 2016	Review Article of SARS-CoV-1, MERS, and Influenza studies	Acute care patient rooms for patients with respiratory infections	Substantial variation in virus survival on surfaces. Temperature and humidity affected virus survival.	Presence of body fluids, such as mucus, substantially increasing virus survival for some studies.
Boone 2007	Mini-review of studies on respiratory virus	Studies on role of fornites in respiratory infections	The authors comment that in many respiratory infections, transmission occurs asymptomatically and multiple routes of transmission may be involved, making it hard to study.	The authors comment that there is strong epidemiological evidence and interventional studies demonstrating the impact of surfaces on transmission of respiratory viruses.
Aiken 2001	Review of studies	Studies on nosocomial spread of viral disease.	Some evidence of surface transmission for a number of respiratory viruses	Interventional measures to stop transmission of respiratory viruses are largely the same, regardless of the virus.

<u>Transmission to a Susceptible Host</u>. Having shown that respiratory virus contamination is common on environmental surfaces, a number of studies have investigated the potential for transmission to hands and from hands to their face.

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Table 4: Select Studies InvestigatingTransfer from Surfaces to Hands andTransmission to a Susceptible Host

Reference	Study Location	Sample Population or Study Environment	Behavior/Activity Observed	Implications or Other Comments
Kwok 2015	University Medical School	Medical students and face touching with hands.	Face touching occurred 23 times per hour, 44% of which involved mucous membrane contact.	When touching a mucous membrane: 36% involved the mouth 31% involved the nose 27% involved eyes 6% were a combination.
Pitol 2021	Review of other studies	Face touching is a common behavior	Face touching occurs 16 – 37 times per hour.	Authors comment the data supports hands being contaminated from environmental surfaces
Rusin 2002	Laboratory study	PRD-1 phage contaminated surfaces	65% of phage transferred from surfaces to hands. 34% of hand contamination could reach the mouth.	Transfer from hard nonporous surfaces was more efficient than from porous surfaces.
Xiao 2018	Acute care facility	Modelling of Influenza outbreak in hospital	Estimated risk for different modes of transmission. Airborne was most likely and fomite transmission was estimated at 6%.	Transmission strongly influenced by virus survival on skin, hand decontamination rates, and contamination of environmental surfaces.
Brankston 2007	Review article	Studies investigating mode of transmission for influenza virus.	No study clearly demonstrated transmission via contaminated surfaces, but substantial epidemiological evidence shows transmission occurs most readily over short distances where surfaces may play a role	3/9 studies, airborne transmission was suggested 6/9 droplet or contact transmission was suggested.
Killingley 2013	Review article	Studies investigating potential for surface transmission of influenza virus.	Strong evidence of environmental survival of influenza A virus on surfaces and hands for periods of time that would allow transmission to occur.	The authors note an absence of direct evidence of transmission by surfaces for influenza.
Winther 2007	Hotel room and laboratory	People with rhinovirus infection	Rhinovirus contamination: 100% of hands at check-in 33% of hands at check-out 35% of environmental sites at check-out.	When surfaces were artificially contaminated with rhinovirus in the lab, touching the surface with hands resulted in transfer in 60% of trials.
Kraay 2018	Modelling study	Surface mediate transmission of norovirus, rhinovirus, and influenza.	Influenza transmission via surfaces most likely in facilities with high hand contact to surfaces.	Risk of transmission reduced by frequent surface disinfection, while hand hygiene 3+ times per day had little impact.

Lastly in this section, we discuss a study by Lei (2017), who modelled the risk of the transmission of several respiratory viruses (SARS-CoV-1 and influenza A H1N1), and norovirus, which is a gastrointestinal/enteric virus, in a Boeing 737 aircraft. While all three viruses were likely to result in transmission when shed by infected people, estimated by type of exposure showed wide variances. While an aircraft interior during flight may not be generalized to the risk of other environmental exposures, this modelling reinforces the idea that mode of transmission is context specific as environmental conditions can play a significant role in transmission risk and some respiratory viruses may be more likely than others to be transmitted via fomites (surfaces).

Table 5. Modelled exposure risk for airplane exposure of Influenza, SARS-CoV-1, and norovirus

	Type and Percent of Estimated Transmission		
Pathogen	Airborne	Droplet	Surface
Influenza	34.3%	65.0%	0.7%
SARS-COV-1	18.2%	20.3%	61.4%
Norovirus	8.1%	0.0%	91.9%

(Source: Lei, 2017)

Interventions that Reduced the Risk of Respiratory Infection

Intervention programs designed to reduce the risk of infection can be designed horizontally or vertically. Vertically designed programs are tailored to reduce colonization, infection, and transmission of a specific pathogen, such as a program to reduce *Escherichia coli* infection, while horizontal programs are designed to implement standard practices and controls that will have an impact on many potential pathogens with a similar mode of transmission with less concern for person specific factors, such as might be found in a program to reduce respiratory virus infection.

While both can have a place in healthcare and public health, horizontal programs are typically favored because they can interrupt the chain of transmission for more pathogens on a more consistent basis. Wenzel (2010) notes that healthcare facilities that attempted

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to reduce the rate of MRSA bloodstream infections from vertical intervention programs did not show a major reduction in rates, but facilities that used horizontal approaches showed larger reductions in rates. Septimus (2014) notes that horizontal and vertical approaches are not mutually exclusive, but horizontal approaches are more cost effective as a general practice while vertical approaches are more appropriate during Gauthier (2020) outbreaks. proposed frequent cleaning and disinfecting within healthcare settings based on care practices, to reduce all microbial pathogens on near patient surfaces.

Several studies in the literature demonstrated a reduction in the risk of respiratory infection with widespread use of horizontal practices, such as surface disinfection, hand hygiene, and good respiratory practices. Three of these studies are discussed briefly.

Table 6: Select Studies Investigating Interventions Targeting Risk Respiratory Illness

Reference	Study	Sample	Behavior/Activity	Implications or Other
	Location	Population or Study Environment	Observed	Comments
Ryan 2001	Military base	Military recruits	Mandatory handwashing 5 times per day and impact on respiratory infection rate	Respiratory infection rate was 45% lower with mandatory handwashing. 49% of recruits reported issues with being able to wash hands, such as lack of time or supplies.
Reynolds 2016	Commercial office	Bacteriophage seeded onto surfaces	Presence of bacteriophage on surfaces was reduced with the intervention of providing facial tissues, hand sanitizer, and disinfectant wipes.	Presence of phage reduced from 47% to 19% for individual workspaces and from 51% to 5% for communal areas.
Chu 2020	Systematic review and meta-analysis	Studies using non- pharmaceutical interventions including distancing, face masks or face shields and eye protection and respiratory virus infection.	Transmission was lower when >1 meter distancing was followed and even lower over longer distances (>2 m). Face masks significantly reduced risk of transmission - stronger risk reduction when N-95 or similar respirators were used versus surgical masks or cotton face coverings. Eye protection was also associated with lower risk or infection	All of the studies reviewed used a bundled intervention, making it difficult to assign a specific reduction in risk to a single intervention and the analysis of the benefit of each intervention was moderate (10-30%), suggesting no single intervention was completely protective.

Modelling Studies Estimating SARS-CoV-2/COVID-19 Risk and Mode of Transmission

Several modelling studies have been published estimating the portion of transmission that occurs by each mode of transmission (contact, droplet, or airborne) or the portion that occurs by the person's symptom status (asymptomatic, presymptomatic, symptomatic). Four of these are discussed below.

Table 7: Select SARS-CoV-2/COVID-19Modelling Studies Investigating Risk ofInfection

Reference	Model goal	Key Model Output	Implications or Other Comments
Ferretti 2020	Estimate portion of transmission by presymptomatic versus asymptomatic versus symptomatic people.	Presymptomatic = 47% Symptomatic = 38% Asymptomatic = 6% Environmentally mediated = 10%	Study done in May 2020 using data available from Jan-Apr 2020. Study predicts that most transmission is by people without symptoms at the time of transmission.
Pitol 2021	Estimate role of surface transmission using Quantitative Microbial Risk Assessment	They estimate the risk of infection from a single hand to surface contact followed by hand to face contact was 0.2% to 5% for the lowest and highest risk scenarios.	This studies used for parameters were confounded by a range of interventions being in place which could influence environmental contamination rates.
Jones 2020	Estimated transmission risk for healthcare workers	Inhalation = 61% Droplet transmission = 32% Contact transmission = 6.9%	Transmission risk was related to the number of hand to face contacts, risk of spray transmission, viral load in saliva, and frequent coughing.
Wilson 2020	Estimate risk of transmission for a single hand to surface contact using Quantitative Microbial Risk Assessment	At low virus level, risk was estimated at 1 in 1 million.	When virus levels on surfaces were 10,000 copies per cm ² and at least 10% of the virus was infective, disinfection would have to reduce virus levels by >2.5 log to achieve the same low level of risk.

Discussion

One of the challenges in using QMRA to assess risk is that while modelling can calculate a level of risk, it cannot determine whether a given level of risk is acceptable. As an example, the risk of contracting HIV from a needle stick, such as might be experienced by a healthcare worker during patient care, is estimated at 0.3% (Singh, 2012) or 1 in 333. Significant workplace safety practices are used in healthcare to protect workers from this risk. If the risk of getting SARS-CoV-2, a disease with an approximately 2% case fatality rate, was similarly <1.0%, one facility might find this

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risk acceptable, while another might find it unacceptable and implement precautions to reduce the risk of infection. Determining what the acceptable level of risk is for a facility is not something that can be determined by modelling.

Context is important in proposing interventions, but the facility context can change over time. Horizontal prevention programs are favored because they help avoid the need to constantly reassess whether your facility context has changed. Surface cleaning and disinfection remain an important intervention for public facilities because it can provide an incremental hygiene benefit across a wide range of facility contexts and this is evident in a wide range of CDC and WHO publications for commercial facilities which encourage cleaning and disinfection of environmental surfaces.

A wide range of pathogens can be transmitted via surfaces including those deposited via respiratory droplets. While the incremental risk from surfaces for SARS-CoV-2, or any specific pathogen, may be low, the composite risk of all pathogenic bacteria, fungi, and viruses that may be present on the surfaces of a commercial facility argue in favor of the importance of regular surface hygiene being a standard facility infection prevention practice.

People who spread pathogens are not always symptomatic. In the case of SARS-CoV-2, it is estimated that ~50% of transmission occurs in asymptomatic or presymptomatic people. Studies of other respiratory viruses show that this can occur for influenza and other respiratory viruses. Physical symptoms such as coughing or sneezing may not indicate a risk of pathogen transmission. allergies and as other conditions may also induce these behaviors, and shedding may occur absent any visible symptoms. A more conservative approach is

to assume that some portion of the people using a public facility are transmitting pathogens and implement control measures to address this risk rather than relying on observation of visible symptoms to trigger using control measures.

Conclusion

This discussed the risk of paper environmental transmission of respiratory viruses and the available evidence that supports the various modes of transmission. While significant data shows the potential for replication competent virus on surfaces in a wide range of settings, data on contact transmission of respiratory viruses is significantly confounded in most studies by the presence of multiple modes of transmission being possible, multiple interventions being used to interrupt transmission, and the preference respiratory viruses have for droplet transmission.

A number of studies have shown that contact transmission is likely to occur in certain sets of circumstances as an is an opportunistic mode of transmission and that frequent hand hygiene and surface disinfection are likely to play a role in reducing this risk when it does occur. Given the wide range of pathogens that can be transmitted via contaminated surfaces, use of frequent surface disinfection as part of a horizontal bundle of interventions to reduce the risk of infection is a sensible precaution.

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