

Supplementary material

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Supplementary Table. Comparison of the four common childhood respiratory viruses¹⁻¹²

	Influenza	COVID-19	Respiratory syncytial virus	Rhinovirus
Epidemiology	<ul style="list-style-type: none"> ➤ Five pandemics in the last 140 years, with the 1918 pandemic being the most severe ➤ Estimated >50 to 100 million deaths ➤ <1 million deaths in the 2009 swine flu pandemic ➤ Pandemics occur irregularly ➤ Aerosol transmission ➤ $R_0 = 1-2$ ➤ Three to four waves of increasing lethality ➤ Mortality was greater at the beginning of a wave 	<ul style="list-style-type: none"> ➤ Repeated zoonoses (eg, SARS, MERS, and now COVID-19) and annual circulation of seasonal coronaviruses ➤ Between 2000 and 2022: >774 million cases confirmed, with >7 million deaths reported ➤ Global death-to-case ratio: 2.1% ➤ Aerosol and airborne transmission ➤ $R_0 = 2-3$ ➤ In waves with containment and mitigation stages 	<ul style="list-style-type: none"> ➤ Most common cause of respiratory hospitalisation in infants ➤ An important pathogen in all age groups ➤ Infection rates higher during the cold winter months ➤ Bronchiolitis in infants, common colds in adults, and more serious respiratory illnesses, such as pneumonia in the elderly and immunocompromised individuals¹ ➤ Outbreaks spread by contaminated air droplets 	<ul style="list-style-type: none"> ➤ High global prevalence² ➤ Infants, the elderly, and immunocompromised individuals most affected³ ➤ Transmission via respiratory aerosols and fomites
Biology of organisms	<ul style="list-style-type: none"> ➤ Enveloped single-stranded RNA virus ➤ <i>Orthomyxoviridae</i> family ➤ Majority influenza A, 	<ul style="list-style-type: none"> ➤ Enveloped single-stranded RNA virus ➤ <i>Coronaviridae</i> family ➤ SARS-CoV-2 ➤ Possible association 	<ul style="list-style-type: none"> ➤ Single-stranded negative-sense RNA virus ➤ <i>Pneumoviridae</i> family ➤ No genetic 	<ul style="list-style-type: none"> ➤ Single-stranded positive-sense RNA genomes ➤ <i>Picornaviridae</i> family

	eg, H1N1, H2N2, H3N2, and H5N1		with antigenic drift		reassortment and antigenic shifts ¹
	➤ Associated with antigenic shift				
New strains/variants	➤ Zoonotic transmission (eg, pigs, chickens, and ducks)	➤	New variants: B.1.1.7, P.1, B.1.351, B.1.427, and B.1.429	➤	Antigenic subtypes A and B
					➤ Three species (A, B, and C)
Pathophysiology of clinical presentation	➤ Haemagglutinin protein of the influenza virus binds to the sialosaccharides of respiratory epithelial cells	➤	Spike protein of the SARS-CoV-2 virus binds to the angiotensin-converting enzyme 2 receptor of olfactory and respiratory epithelial cells		N/A
	➤ Viral replication within the nucleus	➤	Viral replication in the cytoplasm		N/A
Diagnosis	➤ RT-PCR test to detect influenza RNA from upper respiratory tract samples, eg, nasopharyngeal swab, nasopharyngeal aspirate	➤	RT-PCR test to detect SARS-CoV-2 RNA from upper respiratory tract samples, eg, nasopharyngeal swab, nasopharyngeal aspirate, and deep throat saliva	➤	Antigen, molecular testing
		➤	Serologic test to identify previous or late infections	➤	Viral culture
Variable	➤ ~5%	➤	Overall 2% to 3%	➤	At least one RSV
				➤	Not usually associated

mortality	<ul style="list-style-type: none"> ➤ Lower mortality than RSV in children⁴ 	<ul style="list-style-type: none"> ➤ Low mortality; children may be asymptomatic or silent carriers 	<p>infection by the age of 2 years</p> <ul style="list-style-type: none"> ➤ Reinfection is common ➤ More serious infections in 15% to 50% of cases in children, 25% of cases in adults ➤ High mortality in infants and the elderly^{4,5} ➤ 80% mortality in immunocompromised individuals 	<p>with mortality</p> <ul style="list-style-type: none"> ➤ Infants, the elderly, and immunocompromised individuals are most affected
Prevention strategies	<ul style="list-style-type: none"> ➤ Culling and vaccinating livestock ➤ Vaccinating poultry workers against common strains ➤ Limiting travel in pandemic areas ➤ Strategies to slow down a pandemic include public response measures, social distancing, respiratory hygiene, handwashing hygiene, masks, and risk 	<ul style="list-style-type: none"> ➤ Staying at home, universal mask wearing, avoiding crowded places, social distancing, ventilating indoor spaces, thorough hand washing, practising respiratory hygiene, and avoiding touching the eyes, nose, or mouth with unwashed hands ➤ Vigilant contact tracing 	<ul style="list-style-type: none"> ➤ Thorough hand washing, avoiding close contact ➤ Palivizumab in high-risk infants^{5,6} 	<ul style="list-style-type: none"> ➤ Droplet precautions ➤ Thorough hand washing ➤ Surgical mask and gloves

	communication	➤ Travel restrictions and quarantine measures		
		➤ Regular testing		
Antiviral drugs	➤ Oseltamivir and zanamivir	➤ No specific effective antiviral treatment or cure (no good evidence for lopinavir, ritonavir, or remdesivir)	➤ Ribavirin controversial ⁷	➤ Pleconaril is not currently available
	➤ Adamantanes (amantadine and rimantadine)	➤ At risk groups with mild to moderate symptoms can take nirmatrelvir/ritonavir		
		➤ Glucocorticoid (dexamethasone) effective for severe cases		
Vaccines	➤ Several	➤ >13 vaccines	➤ No vaccine yet ⁷	➤ No vaccine yet
	➤ Variable efficacies and side-effects	➤ Variable efficacies and side-effects		
	➤ Vaccine hesitancy	➤ Vaccine hesitancy		

Abbreviations: COVID-19 = coronavirus disease 2019; MERS = Middle East respiratory syndrome; N/A = not available; RT-PCR = reverse transcriptase–polymerase chain reaction; R₀ = reproduction number; SARS = severe acute respiratory syndrome; SARS-CoV-2 = severe acute respiratory syndrome coronavirus 2

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