### What We've Learned So Far:

Identification, Management and Impact of Covid-19 on Chronic Lung Disease

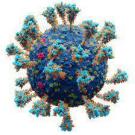


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Nancy Nathenson, RRT Respiratory Care/Health Educator & Consultant Community Asthma& COPD Expert -Allergy & Asthma Network

## Coronaviruses

- First identified a human coronavirus in 1965
- It caused a common cold
- Later that decade, a group of similar *human* and animal viruses were found and were named coronavirus after their "crown-like appearance"
- Primarily cause infections in birds and mammals but have shown to be capable of infecting humans



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## Severe Acute Respiratory Syndrome (SARS)

- China's Guangdong province
- Horseshoe bats-natural reservoir of SARS-CoV
- Spread by civets and other animals in wet markets
- 2003 SARS atypical pneumonia
- 1/3 fever, dyspnea, cough, watery diarrhea and needed mechanical ventilation
- Nov 2002-July 2003 -WHO reported 8,098 SARS cases from 29 countries,
- 774 SARS-related deaths (case-mortality rate: 9.6%)
- 29 cases in U.S. with "0" deaths

# Middle East Respiratory Syndrome (MERS-CoV)

- Demonstrated the lethality of CoV's when they cross the species barrier and infect humans
- First known cases appeared in Jordan in April 2012.
- Largest outbreak 2015 in the Republic of Korea
- WHO reported > 2400 cases linked to the Arabian Peninsula
- Transmitted by camels
- Similar symptoms to SARS but 50-89% required mechanical ventilation
- 35% Mortality







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# Severe Acute Respiratory Syndrome SARS CoV-2

- Term COVID-19- CoronaVIrusDisease-19
- SARS-CoV-2 due to similarity to SARS-CoV
- 1<sup>st</sup> case- December 12, 2019 in Wuhan, China
- Declared a pandemic (WHO) on March 11<sup>th</sup> 2020
- Variety of NEW symptoms dyspnea, neurological disturbances, nausea, headache, vomiting

# SARS CoV-2 Mortality

- 65 years and > = 80% of Covid deaths
- Residents in LTC < 1% of the population-35% of ALL deaths</li>
- Fatalities 10-fold higher in US high population counties
- Racial and ethnic minority groups die at younger ages
- Highest % of increases in mortality among adults 25-44 years of age Hispanic or Latino
- Older adults and males higher mortality risk





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### COVID-19 Contributions to Worsening Outcomes

- Co-morbidities
  - Cardiovascular Disease
  - Diabetes

- Liver DiseaseObesity
- Chronic Pulmonary Disease
   Genetics
- Kidney Disease
- Immune Disorders



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- Racial and Socioeconomic disparities
- Population Density



## Transmission SARS/MERS vs. SARS CoV-2

#### SARS-MERS

- Transmission from recognized disease (symptomatic)
- Grows on standard tissue epithelial cells in the <u>lower</u> lung
   SARS CoV 2



- Transmission from asymptomatic, mildly symptomatic and super spreading
- Frequently transmitted 1-2 days prior to symptoms when most contagious
- Complicated the identification of cases and prevention of spread
- Grows better on epithelial cells of the primary or <u>upper</u> airway
- Upper airway continuously exposed to pathogens and pollutants

### Angiotensin Converting Enzyme-2 (ACE-2)

SARS CoV-2- 10 x's higher affinity to bind with ACE 2 than SARS

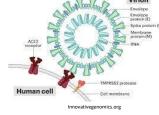
Limiting factor in initial infection

ACE-2 expression varies and affected by chronic inflammatory diseases, Asthma & COPD

ACE-2 increased in COPD and Tobaccos Smokers

Inhaled corticosteroids (ICS) used to treat AR, Chronic Rhinosinusitis, Asthma and COPD

ICS lowers expression of ACE-2

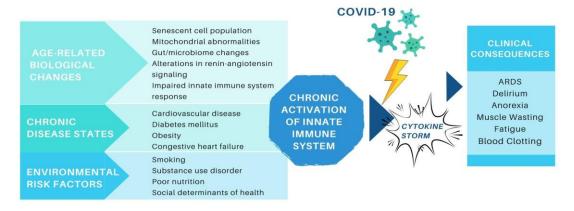


- Biologics upregulate ACE-2 and increase risk of SARS CoV-2
- ACE-2 Inhibiting agents can prevent the viral infection in vitro

## The Aging Immune System

#### **Covid Related Contributions to Cytokine Storm**

And Observable Clinical Consequences



## COPD

≈384 million people world wide
2<sup>nd</sup> most common cause of
disability
What was the 3<sup>rd</sup> leading cause of
death in 2020 and 2021?



### COPD and COVID-19 - Risk & Severity

COPD patients NOT at increased risk of infection from Covid-19 Why?

- Fragile, more careful and utilized prevention measures
- Decreased air pollution due to the lockdown
- If ICS use- suppresses virus replication & cytokine production

#### Comorbid COPD and CLD In COVID-19

- Higher severity, complications & mortality
- Chronic inflammatory state and low respiratory capacity
- Poor lung function reserves, pnx, ARDS, Pulmonary Vascular Thrombolytic events lead to resp failure
- ACE-2 receptors are upregulated in the small airway epithelium and alveoli

# Challenges in Asthma & COPD Dx and Monitoring

Reductions in face to face consultations, rehab and exercise programs and home care

- Overlapping symptoms
- Access to care due to COVID patients
- Disruption of global supply chain
- Inability to afford medications with economic hardships
- Exacerbation events expose individual to increased risk
- Inability to assess inhaler technique
- Infection risks with certain meds
- Lack of telemedicine/Technology unavailable to the patient

## Challenges in COPD Care in COVID-19

- Distinct difference in COPD Exacerbation w/COVID
  - Vascular injury
  - Pneumonitis
  - Coagulopathy
  - Systemic inflammation
  - Cytokine storm
  - Multiorgan involvement

# COPD-Maintenance & Prevention in COVID-19 Pandemic

- If stable remain on ICS and OCS
- Ensure enough medication
- Remain on ACE inhibitors and Angiotensin Receptive Blockers
- Inhalers vs aerosols
- Vaccinations: Pneumoccocal PPSV23,Tdap vaccine
- Prioritize for Covid Vaccination/Boosters
- General strategies, sheltering in place, hand washing, masking but <u>not lead to social isolation</u>
- Stay connected



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# Symptoms: COPD Exacerbation vs. COVID-19

COPD Exacerbation: More coughing than usual,

**SOB**, fatigue , edema, trouble sleeping, changes in sputum

COPD & COVID-19 Overlap: Cough and SOB

**COVID-19:** *<u>Fever, loss of smell or taste</u>*, headaches, lymphopenia, GI symptoms, profound fatigue, sore throat, congestion

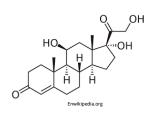
Treat aggressively with systemic corticosteroids and other meds to reduce morbidity and mortality

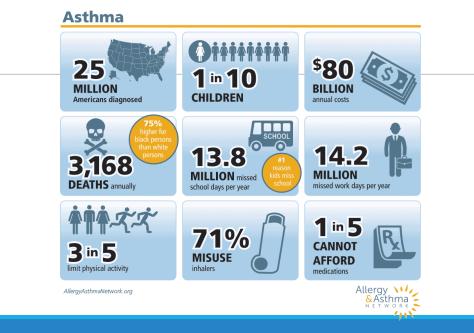


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### COPD w/COVID-19 Special Management Considerations

- Dexamethasone for up to 10 days reduced mortality in patients on Invasive Mechanical Ventilation (IMV) or oxygen alone
- Systemic Glucocorticoids reduce mortality in those NOT on IMV or vasopressors
- Aggressive EXTRA thromboprophylaxis 2 times daily
- Antibiotics with purulent sputum or (IMV)
- At risk for nutritional deficiency and skeletal muscle loss
- Dietary support and early mobilization





## ASTHMA-Dx, Risk & Severity in COVID-19

#### Dx-

- Pulmonary Function Testing, History, Environment, Symptoms
- Spirometry if urgent or essential, Dx or assess lung function prior to surgery, RT-PCR test
- Home Peak Flow measurement, and validated questionnaires ie; Asthma Control Test

#### **Risk & Severity**

Asthmatics-decreased risk of contracting Covid-19

- Decreased ACE-2 expression may lower the risk of Covid 19 severity and mortality
- T helper-2 (Th2) immune response
- ICS use may reduce the risk of infection or developing symptoms

Persons with Moderate to severe asthma are at increased risk of severe illness in COVID-19

# Asthma Maintenance and Monitoring in COVID-19 Pandemic

#### Maintenance

- Continue therapy as usual including Biologics
- Continue ICS
- Little benefit of ICS in acute phases of virus if not already used
- Rescue pack of steroids at home
- Prednisone only for severe exacerbations

#### Monitoring

- Follow Asthma Action Plan
- Peak Flow Diaries
- Asthma Control Tests
- Avoid Triggers
- Review inhaler technique
- Telehealth found to significantly improve asthma control
- Most beneficial in severe disease vs. mild



## Symptoms: ASTHMA vs. COVID-19

#### **ASTHMA SYMPTOMS**

- No fever
- Audible Cough or wheeze
- **Shortness of Breath**
- Chest Tightness and/or congestion
- You have to stop talking to catch breath
- Fatigue
- Not able to perform daily activities

#### SYMPTOMS COVID-19

- Shortness of breath
- Coughing with or without wheeze
- Fever of over 100.4
- Loss of taste or smell
- Headache
- Profound fatigue
- GI Symptoms
- Nasal congestion
- Sore throat

### ALLERGIES Dx, Risk in COVID-19

#### DX:

Personal and Medical History Physical Exam

Testing

- Skin Prick
- Blood tests
- Oral challenges
- Patch

**RISK:** 



There is no evidence to suggest that people with allergies are at greater risk for COVID-19

## Symptoms: ALLERGIES vs. COVID-19

SYMPTOMS: SEASONAL ALLERGIES

- Runny nose, usually with clear or palecolored mucus
- Sneezing
- Coughing
- Red, watery eyes
- Itching around the nose, mouth, or eyes

Symptoms if left untreated

- Shortness of breath
- Nasal congestion, postnasal drip
- Headache
- Sore throat
- Decreased sense of smell
- Sinus, ear infections
- Puffiness or dark circles under the eyes
- Fatigue

- SYMPTOMS: COVID 19
- Fever or chills
- Coughing
- Shortness of breath
- Fatigue
- Muscle, body aches, headache
- NEW loss of taste or smell
- Sore throat, congestion, runny nose
- Nausea or vomiting, diarrhea

# ALLERGY Management & Prevention in COVID-19

- Continue prevention measures Intranasal same for AR, prevent sneezing
- GOLD guidelines-continue biologics
- Take allergy medication 2 weeks before symptoms occur
- General strategies, sheltering in place, hand washing, masking

but not lead to social isolation

- Get the COVID-19 vaccine
- Pfizer/BioNTech and Moderna contain polyethylene glycol (PEG) or polysorbate, may cause allergic reactions in some people
  - Health professionals monitor patients for adverse symptoms for 30min
  - Facilities should be equipped to treat severe allergic reactions
  - Bring your own epinephrine auto-injector for extra protection

# **REMDESIVIR** use in COVID-19

#### REMDESIVIR

Inhibits viral RNA dependent RNA polymerase which inhibits SARS CoV-2 in vitro

- Use for 5 days or until discharge for patients with supplemental O<sub>2</sub> with NO need for HFNC, NIV, IMV or **ECMO**
- EARLIER the administration the greater benefit
  - Decreased LOS reduced recovery time by 4 days most effective in those receiving supplemental oxygen
  - Reduced disease progression
  - Decrease use of respiratory assist devices and ECMO



Monoclonal Antibodies

SARS CoV-2 enters the system through ACE-2 receptors in lungs and GI tract

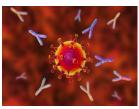
Initially used on compassionate basis w/regulatory approval

In **Low** antibody titer or later in disease course or both:

- Lack of survival benefit
- Does not stop disease progression or help stabilize symptoms
- In High antibody titer/early w/ mild & mild to mod disease
   Slowed progression and better survival
   Decreased progression to hospital and death with IV route
- \*\*\*Recommend for outpatients @ high risk of severe Covid-19\*\*\*

#### **Challenges:**

- Limited availability and delays in administration
- New Variants will render these ineffective .
- Developing Sub Q, IM or inhalation and improving access will impact outpatient care



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# Glutocorticoids

#### Dexamethasone

 RECOVERY TRIAL study showed a GREATER mortality benefit in patients with SEVERE illness with supplemental O<sub>2</sub> and IMV

- When used as a Standard of care
- LESS beneficial in patients with supplemental O<sub>2</sub>
- NO Benefit with patients on Room Air
- Immunosuppressive agents increase viral replication in early illness



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# Medication Nebulization

- Inhalers vs aerosols
- Viral filters
- Mesh nebulizers separate meds from patient interface including circuits
- Proper hygiene and barrier precaution



Presenter image

# Hypoxia/Hypoxemia

Hypoxia- reduced O2 at the tissue level

Anaerobic metabolism with Lactate production

**Hypoxemia**- Drop in the arterial pressure of O2 below 80mmHg

Five causes of Hypoxemia

- Alveolar hypoventilation,
- RT to LT shunt
- Diffusion-perfusion abnormality
- Diffusion impairment
- Ventilation Perfusion mismatch

#### **These result in SOB**



#### "Silent" or "Happy" Hypoxemia

- \* Well tolerated
- \* Early stages on infection

\* No sensation of dyspnea, SOB or increased Work of Breathing

### Non Vent Adults with Hypoxemic Respiratory Failure (HRF)

Begin with Nasal Cannula

- High Flow Nasal Cannula (HFNC) preferred over NIPPV
- HFNC > ventilator free days than conventional O<sub>2</sub> or NIPPV
- 90 day mortality lower in HFNC than conventional O<sub>2</sub> and NIPPV
- HFNC reduced the rate of intubation and ICU mortality

\*\*HFNC and NIPPV use may avoid 20-25% of intubations decreasing use of critical resources during the pandemic\*\*

### Awake Prone Positioning –Non Intubated

- TRIAL of awake prone positioning in Persistent Hypoxemia despite O<sub>2</sub> and NO indication for Invasive Mechanical Ventilation(IMV):
- Awake patients that can adjust position and tolerate prone
- Acceptable in pregnant patients:
  - Lateral decubitus or fully prone
- Contraindications: patients in respiratory distress or require intubation and IMV, hemodynamically unstable, abdominal surgery and unstable spine
- Recommendation AGAINST prone positioning as RESCUE measure for refractory hypoxemia to avoid intubation

### Invasive Mechanical Ventilation (IMV)

Previously intubated early

- When O2 needs exceeded 5-6 lpm
- Fear of spreading the virus
- Closed ventilatory system

Early intubation placed a heavy burden on critical care resources

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Ventilator shortages

2020 studies from Italy and New York recognized a higher mortality with the intubated mechanically ventilated patient

### Impact of Prolonged Mechanical Ventilation

- Delirium
- Hemodynamic instability
- Secondary to decreased sympathetic drive and positive pressure ventilation
- Increased risk of infection
- Immobilization with increase risk of thromboembolism



- Neuromuscular paralysis
- Post ICU syndrome (PICS) physical and neurocognitive dysfunction

### Patient Self Inflicted Lung Injury (P-SILI)

#### Patient Self-Inflicted Lung Injury (P-SILI) in the nonintubated patient with Acute HRF

- High Min. Ventilation, High RR, High Respiratory Drive
- Worsens any pre existing lung injury w/ increased vascular permeability with local and global lung over distension
- Compares to Ventilator-Induced Lung Injury (VILI) caused by high pleural and trans pulmonary pressure swings

# RESCUE Therapies in IMV, Severe Hypoxemia in COVID-19

# Severe Hypoxemia despite optimal ventilation and other strategies

Use recruitment maneuvers , lung protective strategies

 Reduce mortality and improved oxygenation 24 hrs after the maneuver and decreased need for rescue therapy

IF decompensation occurs-DC

- Traditional PEEP strategies decrease mortality
- AGAINST incremental PEEP staircase



Neuromuscular Blockade in COVID-19 IMV Adults w/Moderate to Severe ARDS

Neuromuscular Blocking Agents (NMBA) Intermittent boluses PRN or continuous Continuous NMBA infusion 48 hours

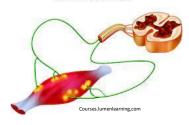
Lontinuous NIVIBA Infusion 48 nours

- In patient ventilator desynchrony
- Ongoing deep sedation prone Ventilations
- Persistent high plateau pressures

#### Monitoring and controlling pain and anxiety

Succinylcholine can lead to hyperkalemia, malignant hypothermia, myalgia, increased gastric intraocular and cranial pressure, cardiac dysrhythmias, (bradycardia) and allergic reactions

Neurons from the spinal cord excite skeletal muscle

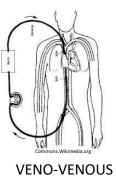


# Extracorporeal Membrane Oxygenation (ECMO)

- Resource intense, nuanced and specialized
- Patient selection, maintenance and liberation
- Indications, Contraindications, Principles remain the same
- Veno-Venous most common for Respiratory Failure with better outcomes
- High Thrombus threat in COVID-19 PTT values of 1.5-2.5 times normal

#### **Extracorporeal Life Support Organization**

40% 90 day mortality rate in 1000 carefully selected patients



# Impact on Chronic Lung Disease Patients in COVID-19

#### ASTHMA in normal times

- Anxiety & Depression
- Mood disorders greater in PWA leads to poor control medication adherence and health outcomes
- Lack of Social support

#### COPD In normal times:

- PTSD, anxiety
- Fear, depression suicidal ideation
- 40% depressive symptoms
- 36% anxiety symptoms



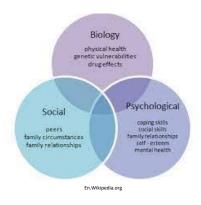
## IMPACT on COPD Patients in COVID-19

- Pandemic may exacerbate worry and symptoms
- Financial stressors bad news, unemployment demoralizing and discouraging
- Social isolation lockdowns feeling trapped hopeless and suicidal ideation
- Bereavement over the loss of loved ones
- Adopt excessive drinking, substance misuse, smoking
- \*\*Important to use Psychological screening tools
- \*\*Depression Anxiety Stress Scale



## BioPsychoSocial IMPACT in Older Adults

"The boundaries between health and disease, between well and sick are far from clear and never will be clear for they are diffused by cultural, social and psychological considerations -George Engel,1977



# **Biological IMPACT**

- Multi organ dysfunction in active and post infection
- NEW physical disability and health deterioration in COVID-19 survivors
- **Respiratory Manifestation**
- ARDS

Neurological

- New disorder
- Or underlying condition



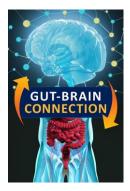
\*\*Cytokine Storm and systemic inflammation responsible\*\*

## Molecular Mechanism IMPACT

Virus infected myeloid and lymphoid cells cause neuroinflammation that linger beyond acute infection leading to neuropsychiatric disorders

Pharmacotherapy

- Immunomodulating corticosteroids
  - Disturbed sleep and cognition
- Gut-Brain Access
  - Alter the microbiome of the gut and brain resulting in psychiatric symptoms



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# Psychological IMPACT

Post Intensive Care Syndrome (PICS)

- PTSD in 30% of ICU admission for ARDS
- High rates of suicide related deaths
- Thwarted belonginess and perceived burdensomeness
- Isolated from family and reliant on care
- Fear of illness, family member burden and disconnection from social contacts
- \*\*\*One study showed a heightened resiliency in those over 60 yrs in those who had strong interpersonal connections\*\*\*

# Social IMPACT

2017 Surgeon General call loneliness a "Global Pandemic" world wide

2018 The UK Appointed a Minister of Loneliness Tracy Crouch

First globally to publish a loneliness reduction strategy

and stated loneliness to be "the greatest public health threat of our time"

Social Isolation increased risk of:

- Heart disease
- Cognitive impairments

Anxiety

- Alzheimers
- Depression
- Death





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# LONG COVID-Post COVID

Symptoms: Fatigue, anxiety, myalgia, low mood and sleep disturbances Risk: No relationship between the severity of the illness and risk of developing Long COVID

#### Findings: Absence of classic respiratory symptoms

- Biopsychosocial effects contribute to long lasting physical and mental symptoms
- Patients with no previous disability are deficient in physical performance

\*\*Prioritization of rehab and psych support and NOT diagnostics and respiratory services\*\*

# Mental Health Impacts on Patient, Family and Healthcare Workers in COVID-19

Admission to the ICU is traumatic for the patient and family

Family involvement in normal circumstances is encouraged and has a positive effect

Families unable to be physically and emotionally there for their loved one

Patients will to live is diminished

WHO speculates this isolation will increase loneliness, harmful alcohol use and drug use, self harm and suicidal behavior

**Healthcare workers** continuously worked in stressful, fearful, resource depleted environments, along with the constant threat of exposure to the virus

Lack of mental health professionals, counselors and programs present an ongoing challenge

# Socio-Economic IMPACT

Income/Job insecurity as a result of decreased global trade

Decreased financial security and reduced earnings

Some forced to retire early:

- Unsafe in workplace
- Unable to telework
- Ageism



**BEFORE** pandemic older adults suffered from food insecurity income based and accessibility barriers

Pandemic created market shortages

Food deserts in rural areas

# LONG COVID-Post COVID

UC Davis Post COVID 19 Clinic - Dr. Christian Sandrock Common symptoms Sleep Disturbances, Fatigue, Lethargy

Treatment focused on better living and high quality of life

Sleep through stress reduction, Meditation and Yoga
 Adjust expectations:

Adjust expectations:

- Difficult to progress a few steps forward and then steps back
- Faltering recoveries, Waxing and waning symptoms
- Requires patience



NIH announced grants as part of it's Post Acute Sequelae of SARS-CoV-2 Infection (PASC)

4/1/2023

# LONG COVID-Post COVID

A retrospective cohort study reviewed 273,618 Covid -19 survivors

**Symptoms:** breathlessness, **fatigue**, chest throat pain , headache, abdominal symptoms myalgia, cognitive and anxiety depression

**Findings:** 1 in 3 had 1 or more features of Long COVID Between 3-6 months after diagnosis

2 in 5 patient recorded Long COVID symptoms in 3-6 month period but <u>none in 1-3 month</u>

**Risk:** Higher in patients with severe illness, females and young people White and Non White equally affected

## Chronic Fatigue in Post COVID-19

- A very large number of viruses can trigger ME/CFS
- Chronic Fatigue Syndrome or ME/CFS
- 17 24 million people worldwide
- United States est. 2.5 million
- All ages and races
- More common in women than men
- The Institute of Medicine reports that patients with ME/CFS are more functionally impaired than people with rheumatoid arthritis, diabetes, and hypertension
- Democratic Rep. Jamie Raskin of Maryland is co-sponsoring a bill, <u>HR 7057</u>, the "Understanding Covid-19 Subsets and ME/CFS Act."



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## Healthcare Workers and Post COVID

A study of persistent post COVID symptoms in HCW

- Covid-19 greater lethality in Males
- Females more to suffer from long term COVID
- After 4 months of the peak wave 45% had symptoms persisting and 32% indicated difficulty coping
- 39% Moderate to severe fatigue

Universal symptoms

- Mild to moderate SOB
- Anxiety Sleep disturbances

ONLY 16% consulted GP only 2% taking sick leave after initial recovery

Even with acknowledgement of severity and struggle to cope



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HAND WASHING/ SANATIZING



### <mark>MASKING</mark>

















SOCIAL DISTANCING













Long COVID SOS Patient-Led Research Collaborative HR 7057 www.nih.gov www.hopkinsmedicine.org www.cdc.org www.fda.gov www.allergyasthmanetwork.org



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