Welcome to the Front Lines of the Fight Against COVID-19

## A TOWN HALL CONVERSATION

We will begin at 10 a.m.





# COVID-19 Effect on the Heart & Vessels

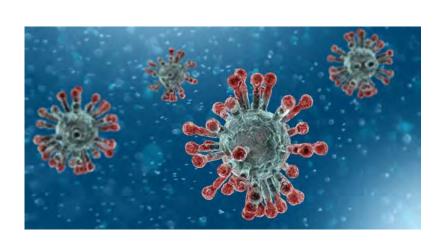
### William A. Zoghbi MD, MACC

Professor and Chairman, Department of Cardiology Elkins Family Distinguished Chair in Cardiac Health Houston Methodist Hospital

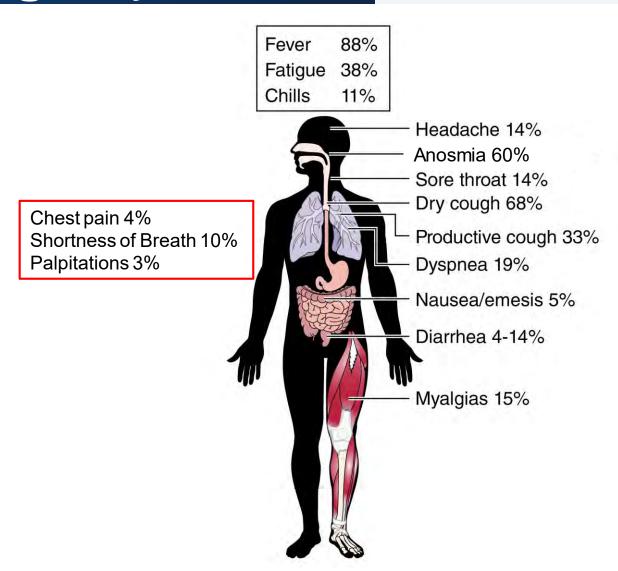


## SARS-CoV-2 Symptoms Involve most Organ Systems



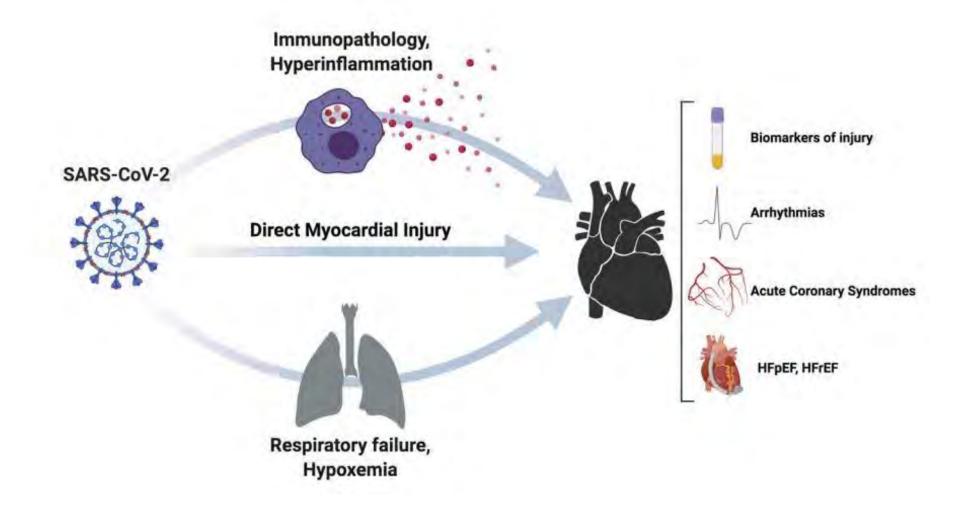


25% Asymptomatic50% Mild symptoms15% Moderate symptoms



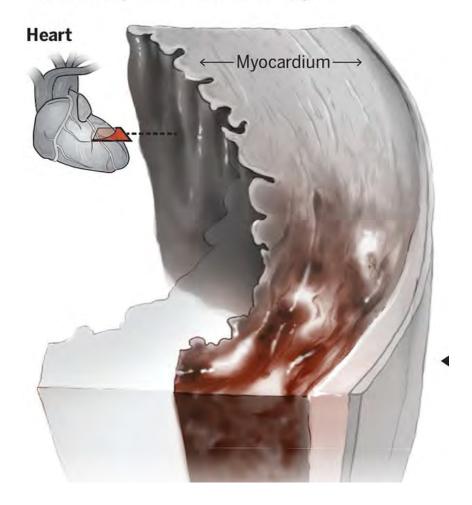
## Cardiac Involvement

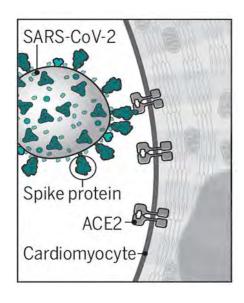


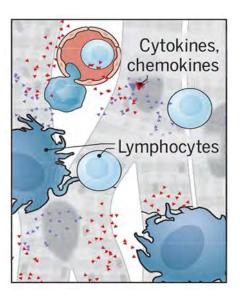


### **Damaging the heart**

Severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection has the potential to directly and indirectly induce cardiac damage.







The Spike protein can bind with ACE2 receptors in vessels throughout the body, enhancing clot formation



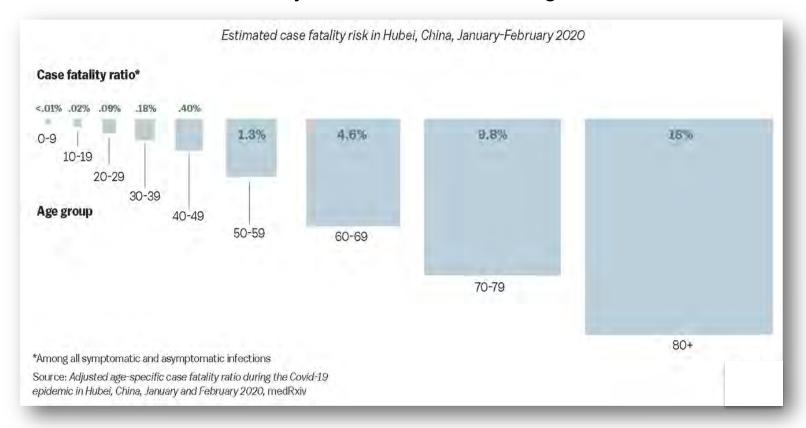
## Risk Factors & Mortality in COVID-19



### Case Fatality Rate Increases with Age

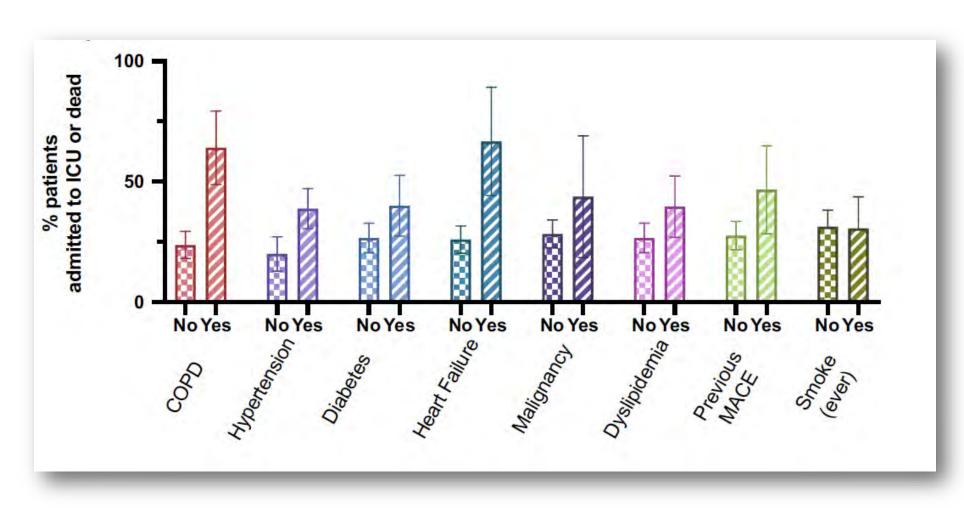
### **Risk Factors:**

- Age
- High Blood pressure
- Diabetes
- Heart Disease
- Obesity
- Lung disease
- Other co-Morbidities



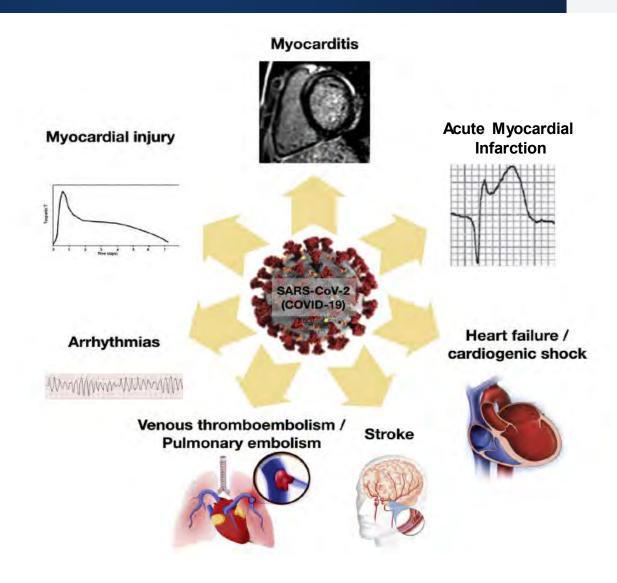
## Risk Factors in COVID-19 Associated with Worse Outcome





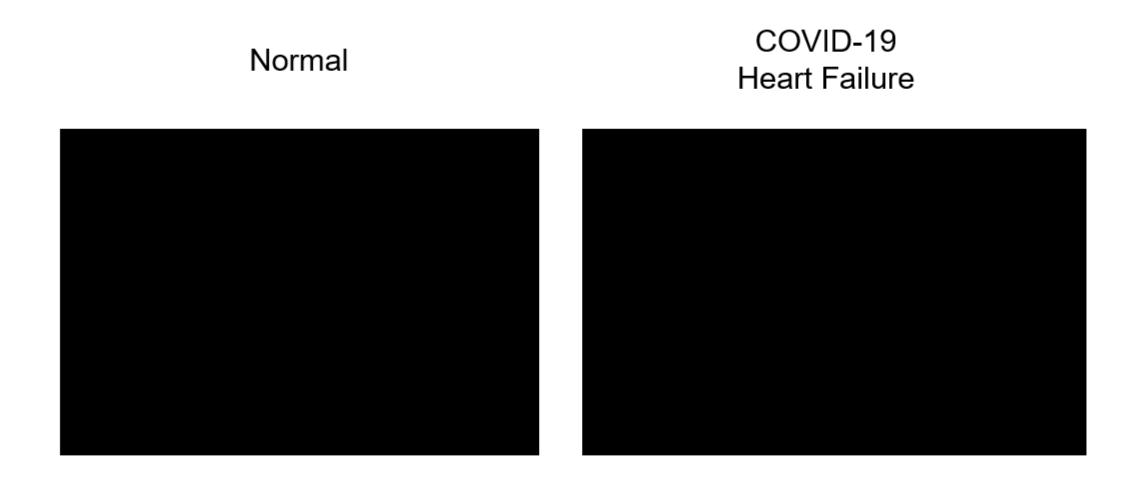
## Involvement of The Heart and Vessels in COVID-19





## Echocardiograms- Heart Function

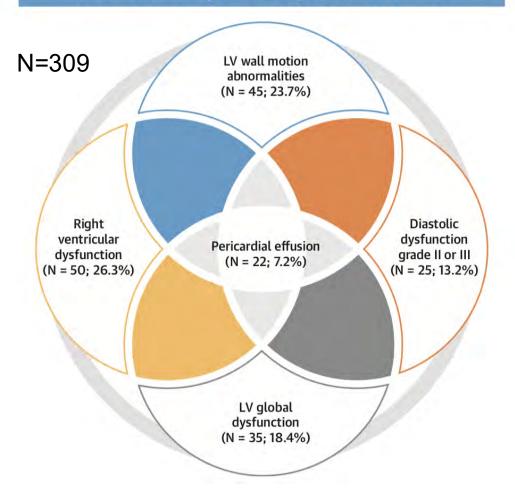


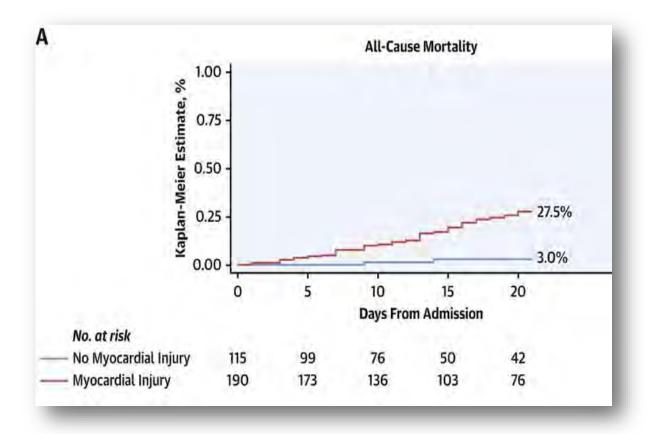


## CARDIAC DYSFUNCTION LEFT vs RIGHT VENTRICLE INVOLVEMENT?



Spectrum of Major Echocardiographic Abnormalities in Patients With Myocardial Injury and COVID-19





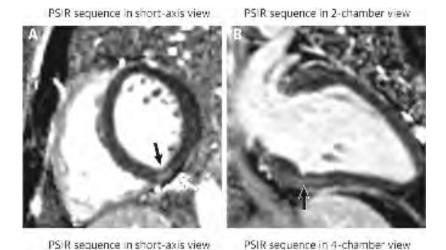
## Cardiac MRI in Recovered COVID-19 Patients



26 patients underwent CMR- median 47 days after COVID-19 symptoms onset

Reported cardiac symptoms (chest pain/palpitations) after recovering from COVID-19

Abnormal CMR- 58% Myocardial edema- 54% Focal fibrosis- Late gadolinium enhancement (LGE)- 31%



## CARDIAC INVOLVEMENT in YOUNG ATHLETES?



JACC: CARDIOVASCULAR IMAGING

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VOL. I, NO.

#### CLINICAL RESEARCH

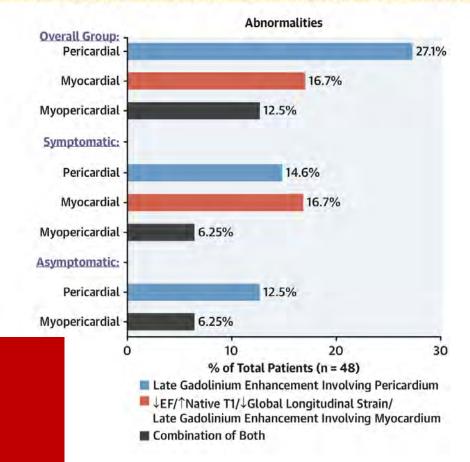
## High Prevalence of Pericardial Involvement in College Student Athletes Recovering From COVID-19

Daniel Brito, MD,<sup>a,\*</sup> Scott Meester, MD,<sup>b,\*</sup> Naveena Yanamala, MS, PhD,<sup>a,\*</sup> Heenaben B. Patel, MBBS,<sup>a</sup> Brenden J. Balcik, MD,<sup>b</sup> Grace Casaclang-Verzosa, MD,<sup>a</sup> Karthik Seetharam, MD,<sup>a</sup> Diego Riveros, MD,<sup>b</sup> Robert James Beto II, MD,<sup>a</sup> Sudarshan Balla, MD,<sup>a</sup> Aaron J. Monseau, MD,<sup>b</sup> Partho P. Sengupta, MD, DM<sup>a</sup>

54 consecutive student athletes with COVID 66% moderate COVID-19 related symptoms 40% athletes with pericardial late enhancements with associated pericardial effusion.

Of the individuals with pericardial enhancements, 6 (12.5%) had reduced global longitudinal strain and/or an increased native T<sub>1</sub>

CENTRAL ILLUSTRATION Abnormal Findings of Pericardial, Myocardial, and Myopericardial Involvement in Mild Symptomatic and Asymptomatic Athletes With Coronavirus Disease 2019

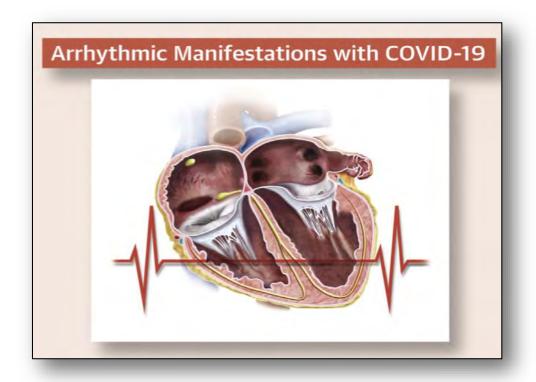


## **ARRHYTHMIAS IN COVID-19**



## Accompany Severity of the Illness & Cardiac Involvement

- Atrial Fibrillation
- Tachycardia
- Heart Block
- Ventricular Tachycardia
- Cardiac Arrest



## ADULT CONGENTIAL HEART DISEASE?



## Moderate/Severe COVID-19 Infection Associated with:

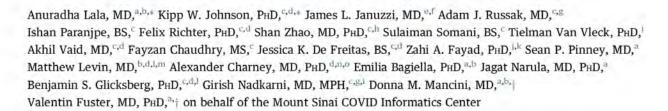
- Genetic Syndrome
- ACHD Physiologic Stage C
- Pulmonary Hypertension
- Obesity

## MYOCARDIAL DAMAGE AMONG COVID PTS PREVALENCE AND RISK



JOURNAL OF THE AMERICAN COLLEGE OF CARDIOLOGY PUBLISHED BY ELSEVIER ON BEHALF OF THE AMERICAN COLLEGE OF CARDIOLOGY FOUNDATION

Prevalence and Impact of
Myocardial Injury in Patients
Hospitalized With COVID-19 Infection

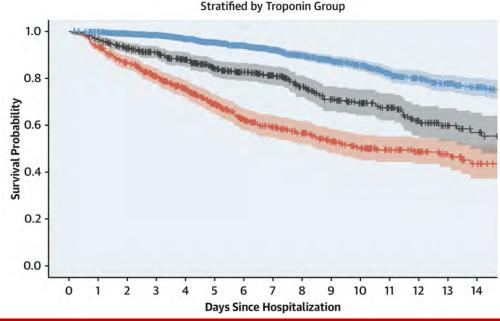


## **2736 COVID patients**

506 (19%) patients died during hospitalization

985 (36%) elevated troponin within 24 hours





Mild Elevation > 0.03-0.09 HR 1.75; 95% CI: 1.37 to 2.24

High Elevation>0.09 ng/dl HR: 3.03; 95% CI: 2.42 to 3.80 What we know about the long term consequences of getting COVID-19



## MAJORITY OF HOSPITALIZED PATIENTS DON'T RECOVER IN 2 MONTHS



## Persistent Symptoms in Patients After Acute COVID-19

JAMA. Published online July 09, 2020. doi:10.1001/jama.2020.12603

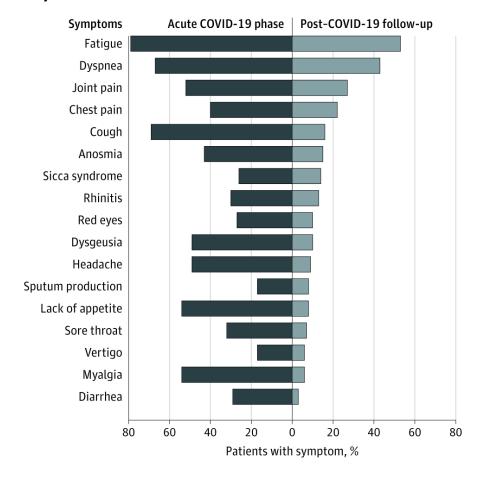
143 Patients Admitted with COVID-19

72% with PNA

2 Week LOS

2 Months post discharge

- 12% completely free of any COVID-19–
  related symptoms
- 32% had 1-2 symptoms
- 55% had 3 or more symptoms



### WHY PROLONGED SYMPTOMS WITH COVID-19?



#### Neurologic

Headaches
Dizziness
Encephalopathy
Guillain-Barré
Ageusia
Myalgia
Anosmia
Stroke

#### Renal

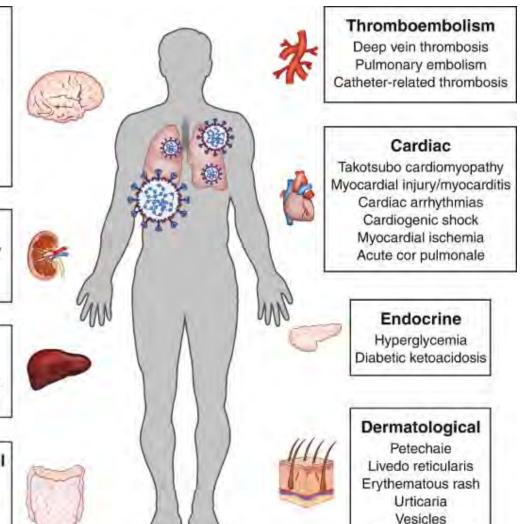
Acute kidney injury Proteinuria Hematuria

#### Hepatic

Elevated aminotransferases Elevated bilirubin

#### Gastrointestinal

Diarrhea Nausea/vomiting Abdominal pain Anorexia

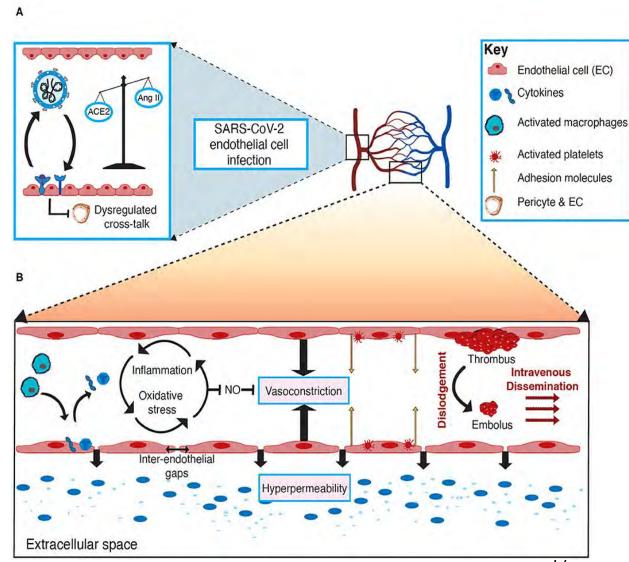


Pernio-like lesions

## Vascular Manifestations-Endothelial dysfunction and thrombosis



- SARS-CoV-2 infects endothelial cells through binding to ACE2 receptor
- Vascular inflammation
- Can lead to vasoconstriction
- Activation of platelets can lead to thrombosis and potential emboli (4-8%)



## COVID-19 Effect on the Heart & Vessels

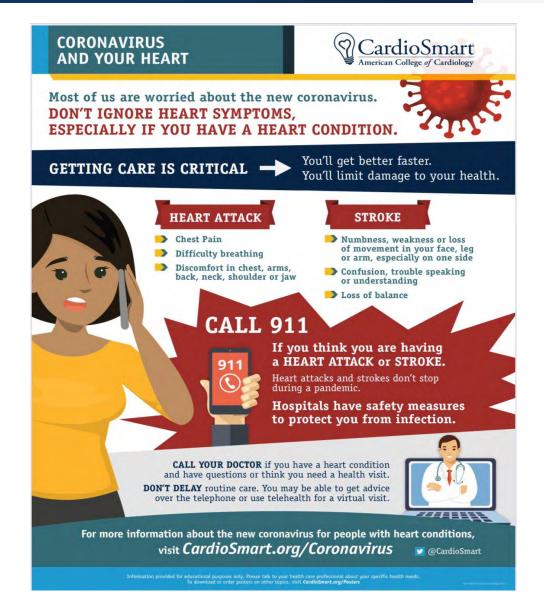


- SARS-CoV-2 can cause heart damage either directly, through an inflammatory process, microthrombi or indirectly because of severe lung damage
- COVID-19 can be associated with myocardial infarction, heart failure, pericarditis or arrhythmias
- Vascular thrombosis (arterial and venous) is seen in COVID-19
- Prognosis is worse in patients where the heart and other organs are more severely affected
- Long term prognosis awaits further observational studies

We are following COVID-19 patients in the CURATOR Registry in the Houston Methodist System with acute and long term comprehensive studies.

## Nationwide Appeal from the American College of Cardiology







## COVID-19 Impact on Surgical Services

Dr. A. Osama Gaber

Chair, Department of Surgery

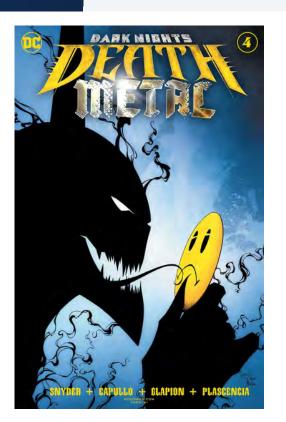
Houston Methodist Hospital



## Surgery Faces the Coronavirus



- In early 2020 the Coronavirus started spreading into the United States
- In March 2020 our community cancelled part of the Houston Rodeo to limit the spread of the disease



## We Faced Amazing Challenges



- Supplies and testing
  - Respiratory supplies
  - Laboratory reagents and supplies
  - Price gouging
  - Disinfectant wipes
- Knowledge gaps
  - Transmission and structure of the virus
  - · Droplet or aerosol
  - Intubation and operation and transmission
- Patient / staff care challenges
  - Delay in diagnosis
  - Virtual care
  - Keeping a healthy staff
- Operational challenges
  - Finances, through put, protection of families

- The Governor's Executive Order
- All licensed health care professionals and all licensed health care facilities shall postpone all surgeries and procedures that are not medically necessary to diagnose or correct a serious medical condition of, or to preserve the life of, a patient who without timely performance of the surgery or procedure would be at risk for serious adverse medical consequences or death, as determined by the patient's physician

## The Long Road to Recovery





You do not have to be great to start, but you have to start to be great

## **HMH SURGERY VISION & MISSION**



#### Vision:

To become the leading academic surgical department in the region

#### Mission:

To lead research and innovation in surgical expertize while providing unparrelled care to our patients, and education for the next generation of surgeons.

#### **Strategic Priorities**

#### Organizational Excellence

- Evolve care delivery into centers rather than practices
- Integrate surgical services across system
- Expand service offerings
- Re-examine departmental resources and revenue streams

#### Education & Professional Development

- Integrate residents/fellows into department productivity
- Capitalize on EnMed
- Emphasize faculty development
- Explore expanding educational offerings in MITIE

### Discovery & Research

- Create a division of surgical sciences
- Increase research funding
- Recruit research faculty
- Enhance research opportunities within department

#### Clinical Excellence

- Grow robotic surgery
- Recruit and retain clinical faculty
- Optimize APP utilization
- Enhance Patient experience
- Create competitive advantage for centers of excellence

#### Collaboration & Community Integration

- Establish deeper collaboration with non-PO physicians and programs
- Grow
   philanthropic
   support
- Enhance department's national and international reputation



We as surgeons share the belief that all human life is sacred. We are devoted to saving, preserving, and improving life regardless of race, color, creed, gender, sexual orientation, and/or place of origin. We also embrace diversity in education, research, employment and every other human endeavor.

Racism needs to be recognized & eradicated.

## DEPARTMENT ASSETS, SPECIALTIES AND FACULTY



#### Bariatrics/MIS

- Vadim Sherman
- Nabil Tariq
- Patrick Reardon
- Lee Morris
- · Debra Harvey (PCP)

#### ACS/Critical Care

- Svlvia Martinez
- Daniel Bonville
- Hina Faisal
- Atiya Dhala
- Rodolfo Oviedo
- Nicole Tapia (Dec 7)

#### Thoracic Surgery

- Min Kim
- Ray Chihara
- Edward Chan

#### General Surgery/Surgical Oncology

- Wade Rosenberg
  - Sherry Lim
  - Michael Reader
  - Nestor Esnaola
  - Feibi Zheng
  - Gilchrist Jackson
  - Alexander Stark
  - · Richard Caplan

#### Transplant

- Osama Gaber
- Richard Knight
- Stephanie Yi
- Hemangshu Podder
- Mark Hobeika
- Mark Ghobrial
- Ashish Saharia
- Constance Mobley
- Robert McMillan

#### Plastics

- Jeffrey Friedman
- Tue Dinh
- Michael Klebuc
  - Aldona Spiegel
- Pierre Chevray
   Eugene Alford
- Evan Collins
- Evan Collins
- Amy S. Xue
- Matthew Kaufman
   Lisa Haubert

#### Woodlands

Baytown

Laura Choi

Shawn Tittle

Robert Ochoa

Esther Dubrovsky

Boutros Kahla

Kendall Sowards

· Timothy Opperman

Ali Mahmood

Sandra Templeton

Michael Yaakovian

Michael O'Shea

Kathleen Baxter

Ziad Kronfol

Sugarland

- Steven "Ben" Albright
- Steven Gordon

#### Willowbrook

- Liza Thalheimer
- Todd Worley
- · Diego Carlos Marines
- Anthony Echo
- John Shuck

#### Clearlake

- Kyle Stephens
- David Laron Mason
- Jitesh Joshi
- Gabe Rodriguez

#### West

- SueEllen Arentz
- Carla Braxton
- Minh Nguyen
- Van-Hien Tran
- Laura Choi
- Warren Ellsworth Rodger Brown

#### **HM Affiliated Surgeons**

- Joshua Coursey
- Eric Haas
- Jean Paul Lefave
- Rachel Ellsworth
- Matthew Weaver
- Randolph Bailey
- Marianne Cusick
- Bidhan Das
- Michael Snyder
- Jeffrey Van Eps

#### Research Faculty

- Xian Chang Li
- · Luping Huang
- Malgorzata Kloc Stepkowska
- Omaima Sabek
- Stefano Casarin
- Xiang Xiao
- Bin He
- Jin Wang
- Wenhao Chen
- · Zhiqiang Zhang
- Linda Moore
- Joshua Swan
- Stephen Jones

## ASSETS: QUALITY METRICS

Mortality, LOS and Readmission
May 2020 YTD, April 2020 Observed\*

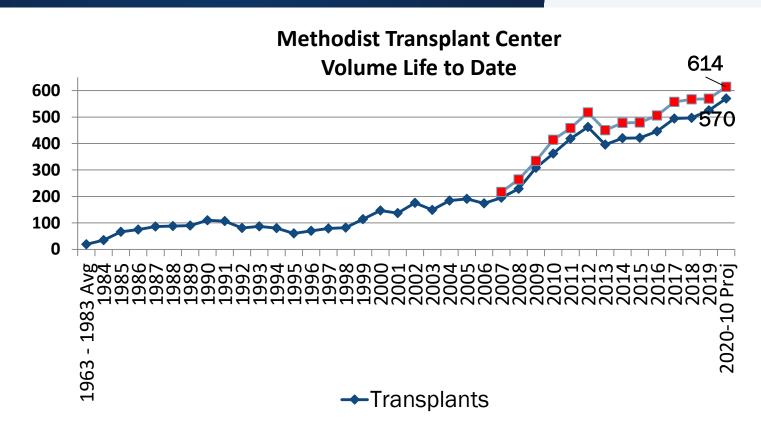


General Surgery Services	2018 3617	2019 3901	2020 YTD May 1162	2020 Goals
Mortality Index	.43 (52)	.43 (56)	.29 (30)	.62 (95 <sup>th</sup> percentile)
LOS Index	.90	.85	.81	.87 (80 <sup>th</sup> percentile)
Readmission	11.67%	11.4%	12.3 %*	10.3% (50 <sup>th</sup> percentile)
Thoracic Surgery Services	2018 223	2019 <sup>299</sup>	2020 YTD May 94	2020 Goals
Mortality Index	0.43 (3)	0.25 (2)	.29 (1)	0.17 (90 <sup>th</sup> percentile)
LOS Index	.78	.87	.75	.82 (80 <sup>th</sup> percentile)
				6.5% 50th

Source: HMH Quality Matrix, May 2020 YTD DW

### **ASSETS: TRANSPLANT**





## We went to work





- Provide top quality surgical care to COVID and non-COVID patients
- Become part of the solution for COVID infections
- Execute on our mission, in care, innovation, education, research and community relations

### 2020 COVID-19 ACCOMPLISHMENTS



### Ongoing Surgery - Transplant - Hepatology Clinical Trials

<u>Title/Topic</u>	<u>PI name</u>	<u>Target/Mechanism of</u> Action	COVID Stage	Inpt or Outpt?
RLF-100 for the Prevention and Treatment of Acute Lung Injury/Acute Respiratory Distress Syndrome in COVID-19 (NeuroRX)	Youssef	Vasoactive Intestinal Polypeptide (VIP)	Severe	Inpatient
Clazakizumab E_IND request: Anti IL6 for use in COVID Patients	Huang	IL-6	Moderate	Inpatient
A Randomized, Double-blind, Placebo- Controlled, Phase 2 Study to Evaluate the Efficacy and Safety of LY3819253 in Participants with Mild to Moderate COVID-19 Illness	Huang	anti-SARS-CoV2 IgG1 antibody	Mild/Moderate	Outpatient
A prospective, controlled, randomized study to compare the efficacy of a chloroquine analog (GNS561) versus standard of care in patients with advanced or metastatic cancer and SARS-CoV-2 (COVID-19) infection		Anti-Viral (PPT-1 modulator/inhibitor)	Mild/Moderate	Outpatient

### 2019 and 2020 METRICS



<u>Grants</u>		Applied for amount or awarded
Grants submitted 2019	39	\$19,467,446.48
Grants awarded 2019	12	\$2,829,123.00
Grants submitted 2020	20	\$17,759,003.98
Grants awarded 2020	10	\$1,577,890.00
Clinical Trials	<u>2019</u>	<u>2020</u>
Surgery	43	49



Payments for Clinical Trials Received to Date 2020: \$660,374.87

### FELLOWSHIP PROGRAMS



#### Surgical Critical Care Fellowship

- Current Status: Initial Accreditation with ACGME
- First fellow graduation in July 2020
- One year program with 2 fellows
- Program Director: Constance Mobley, MD, PhD, FACS

#### Advanced GI/MIS Bariatric Fellowship

- Current Status: Continued Accreditation with MIS Fellowship Council
- Began in 1996
- One year program with 2 fellows
- Program Director: Patrick Reardon, MD, FACS

#### Multi-Organ Abdominal Transplantation Fellowship

- Current Status: Continued Accreditation with ASTS
- Began in 2014
- Two year program with one fellow per year
- Program Director: Osama Gaber, MD, FACS, FAST

### AGCME SURVEY RESULTS



#### 2018-2019 Survey Results (2019-2020 results not yet received)

Evaluation Category	Program Average	National Average
Clinical and Educational Work	4.7	4.8
Faculty	4.3	4.3
Evaluation	4.5	4.5
Educational Content	4.5	4.4
Resources	4.4	4.4
Patient Safety/Teamwork	4.6	4.4

- At or above national average in 5 categories
- 89% of residents reported an overall positive or very positive evaluation of the program

### **ACHIEVEMENTS**



#### **Scholarly Activities**

Increased number of publications

2018 65 publications

2019 70 publications

2020 exceeded 70 publications already

#### **Finances and Revenue**

- Almost back to 2019 activity levels in surgery, and expect to exceed them by the end of December
- Maintaining the budget very close to planned with a small (1-2%) variance

## DEPARTMENT CONTRIBUTIONS TO THE SYSTEM



- Our critical care intensivists worked 24/7 to cover hospital COVID units while our general surgeons with critical care credentials picked up extra shifts to cover the SLICU
- Our ambulatory/outpatient Advanced Practice Providers signed up for extra shifts or transferred to the inpatient units to provide much needed staffing coverage (despite not having worked inpatient units in years)
- Non-clinical staff joined the Labor Pool to assist with screening and other functions while others flexed their hours to allow the department to continue operations, minimizing financial impact

#### FACULTY AWARDS AND ACCOLADES 2020



\*HM System Quality & Patient Safety – Unparalleled Quality Award\*

\*Outstanding Service Faculty
Teaching Award\*
\*I CARE Gold Badge\*

\*ASTS Health Disparities
Research Education Program
(HDREP) Scholar Award\*
\*I CARE Gold Badge\*

\*Exemplary Service Faculty Teaching Award\*







# New Faculty Appointment Mei Rui, PhD





- BA, Yale University
- MM, Yale School of Music
- AD, Yale School of Music
- DMA, Stony Brook University
- Artist in Residence: Yellowbarn, Putney,
   VT
- Research Scientist at Center for Performing Arts Medicine
- MUSICARE support <u>video message and a</u> Mozart duet

# Joshua T. Swan, PharmD, MPH, FCCM, BCPS

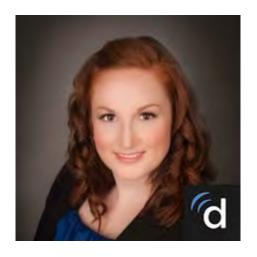




- Associate Professor in Surgery on the Investigative Track
- BS, University of Mississippi
- PharmD, University of Mississippi
- MPH Epidemiology, UT Health Sciences

### Nicole Tapia, MD





- Trauma and Surgical Critical Care Director, HCA Houston Healthcare Kingwood
- BA, UT Austin
- MD, UT San Antonio
- Residency and Critical Care Fellowship, Baylor College of Medicine

# Rodolfo J. Oviedo, MD, FACS, FASMBS





- Fellow of the American
   College of Surgeons
- Fellow of the American
   Society for Metabolic and
   Bariatric Surgery
- Diplomate of the American Board of Surgery

# We Continue to Battle the Coronavirus









## COVID-19 Update

December 10, 2020



# WHAT IS THE CURRENT RATE OF COVID TRANSMISSION IN HOUSTON?

### Houston Area Rt Estimate Trend



#### Rt estimate

This graph shows the R(t) over time. R(t) is a measure of contagiousness or how many people one COVID19 person infects. If R(t)>1, the epidemic is increasing. If R(t)<1, the epidemic is declining. There is higher alert if the whole interval is above the horizontal line at 1. For  $\bf Q$  - Houston, the rate of contagiousness is 1.05; the epidemic is increasing.



### TMC Dashboards





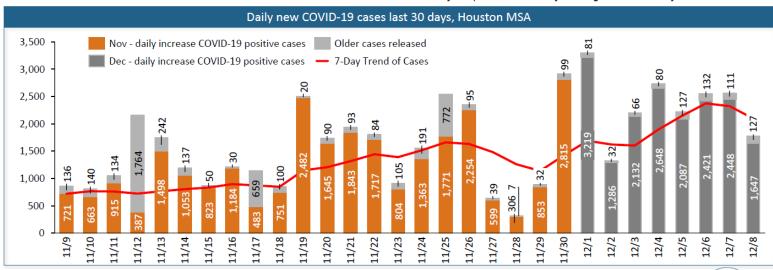
COVID-19 CASE TRENDS

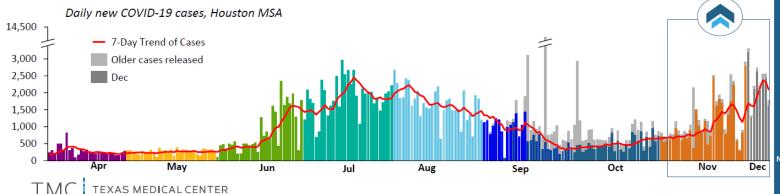
#### DAILY NEW COVID-19 POSITIVE CASES

Greater Houston Area<sup>1</sup>

"TMC" refers to the group of systems that make up Texas Medical Center

Note: On 11/19 Montgomery County batch released cases from previous 2 weeks following database transfer





**December 8, 2020** 

#### **Monitoring threshold:**

Threshold is exceeded by the occurrence of a positive daily growth rate (averaged over 7 days) in the new daily case trend

#### **Current status:**

9 days of positive daily growth rate (averaged over 7 days) in the new daily case trend

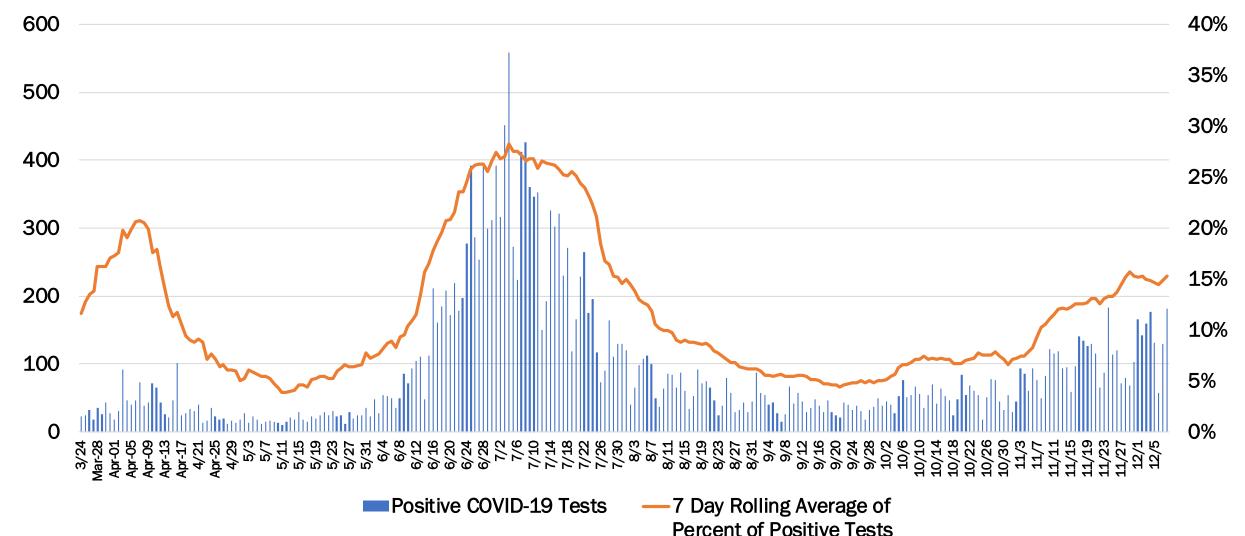
Note: Data for MHHS current as of 12/7 pending update

This document is solely intended to share insights and best practices rather than specific recommendations. 8 Individual institution data is shown as reported and has not been independently verified

### Houston Methodist Testing Trend



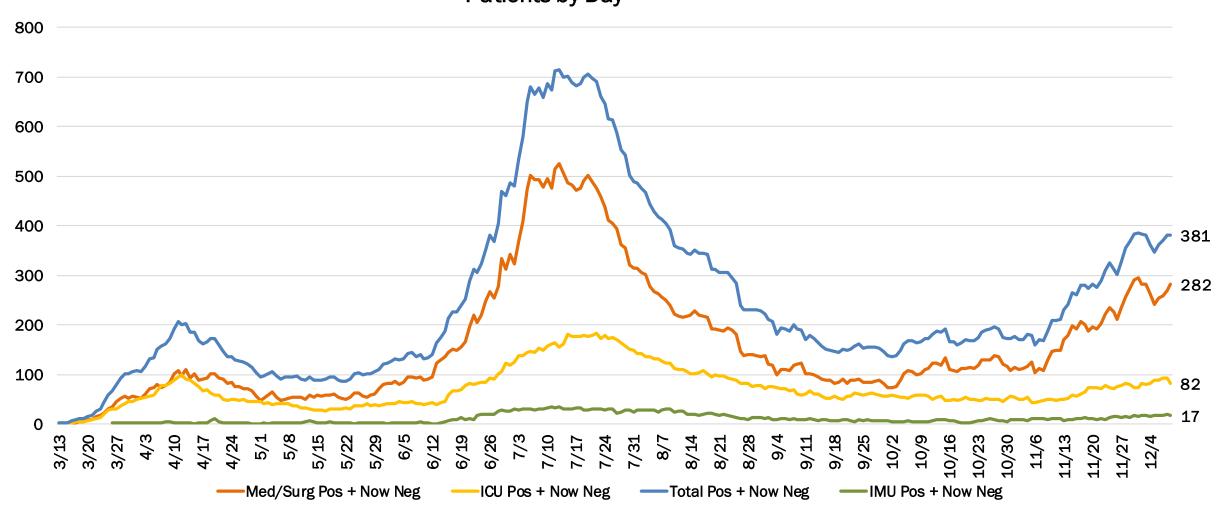
#### Confirmed COVID-19 Lab Tests



# Houston Methodist COVID-19 Cases by Day



### Houston Methodist COVID-19 Patients by Day



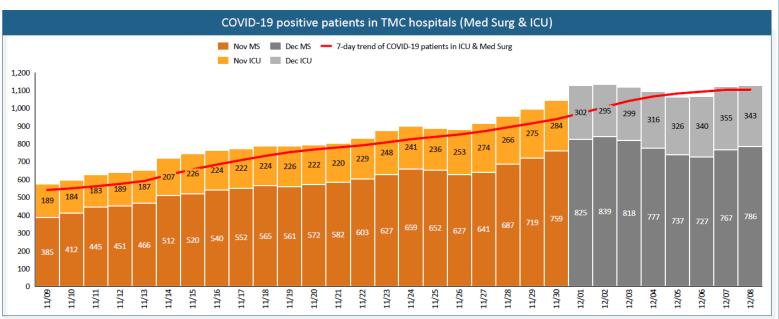
### TMC Dashboards

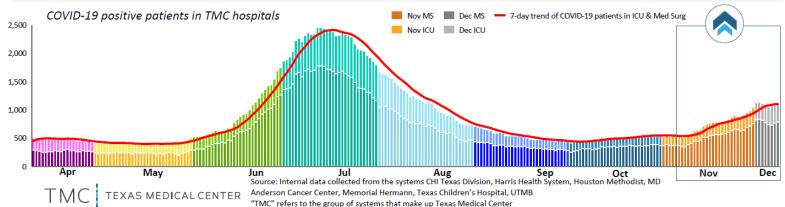


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COVID-19 TMC HOSPITALIZATION

#### TOTAL TMC COVID-19 POSITIVE PATIENTS IN HOSPITAL





All guidelines should be in accordance with CDC guidelines

December 8, 2020

#### **Current status:**

**0.6% total daily growth rate** (averaged over 7 days) in COVID-19 patients TMC hospitals

- 2.5% ICU daily growth rate
- -0.2% Med Surg daily growth rate

#### **Notes:**

While new daily cases may fluctuate for a variety of reasons (e.g., testing), the number of COVID-19 positive patients being treated in med surg and ICU shows an objective view of how COVID-19 impacts hospital systems

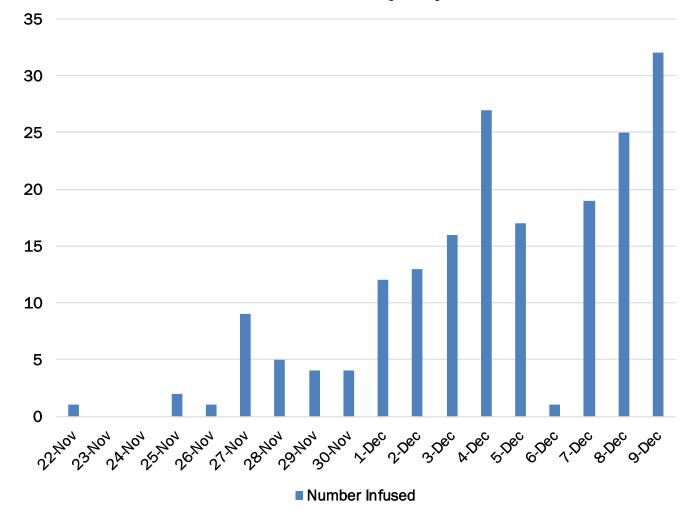
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### Monoclonal Antibody Treatment



### Number of Patients Infused with Monoclonal Antibodies by Day



#### **Emergency Use Authorization Criteria**

- Positive SARS-CoV-2 viral test
- Mild to moderate symptoms
- At increased risk for progression to severe COVID-19 and/or hospitalization
  - BMI > = 35
  - Chronic kidney disease
  - Diabetes
  - Immunosuppressive disease and/or treatment
  - Age>=65
  - Age>=55 AND cardiovascular disease,
     hypertension, and COPD/chronic
     respiratory disease
- Within 10 days of symptom onset

# I HAVE BEEN FOLLOWING SWEDEN IN THE NEWS. WHAT DO YOU THINK ABOUT THEIR APPROACH TO CONTROLLING THE VIRUS?

# Sweden Imposes Mandatory COVID-19 Measures



### THE WALL STREET JOURNAL.

#### Sweden toughens up on COVID-19 restrictions as cases surge

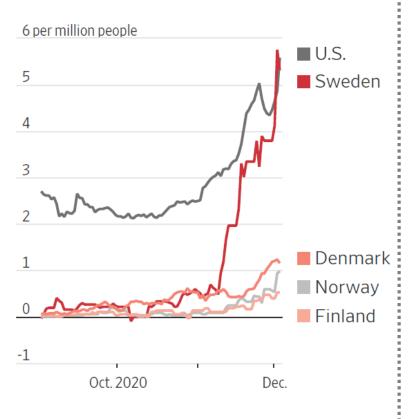
By Jackie Salo

December 6, 2020 | 12:05pm | Updated



Passengers are crammed into a packed subway car in the middle of the ongoing COVID-19 pandemic in Stockholm

### Daily confirmed Covid-19 deaths, seven-day rolling average

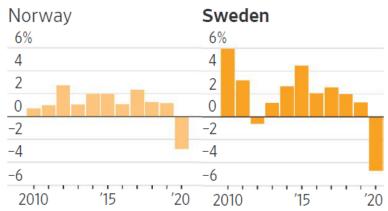


#### **Economic Downturn**

Sweden's gross domestic product and unemployment rate have reached the worst levels in a decade.

#### Percent change in gross domestic product





A NUMBER OF MEDICAL OBSERVERS HAVE SUGGESTED THAT IN FACT THE NUMBER OF INFECTED INDIVIDUALS IS AS MUCH AS 8x THE NUMBER IN THE OFFICIAL COUNTS, GIVEN THE VERY LARGE NUMBER OF ASYMPTOMATIC OR VERY MILD CASES. DO YOU HAVE A VIEW ON THIS? IF IT WERE THE CASE, WHAT ARE THE IMPLICATIONS FOR HOW WE ARE DEALING WITH THE PANDEMIC?

### Estimate of un-Diagnosed COVID-19





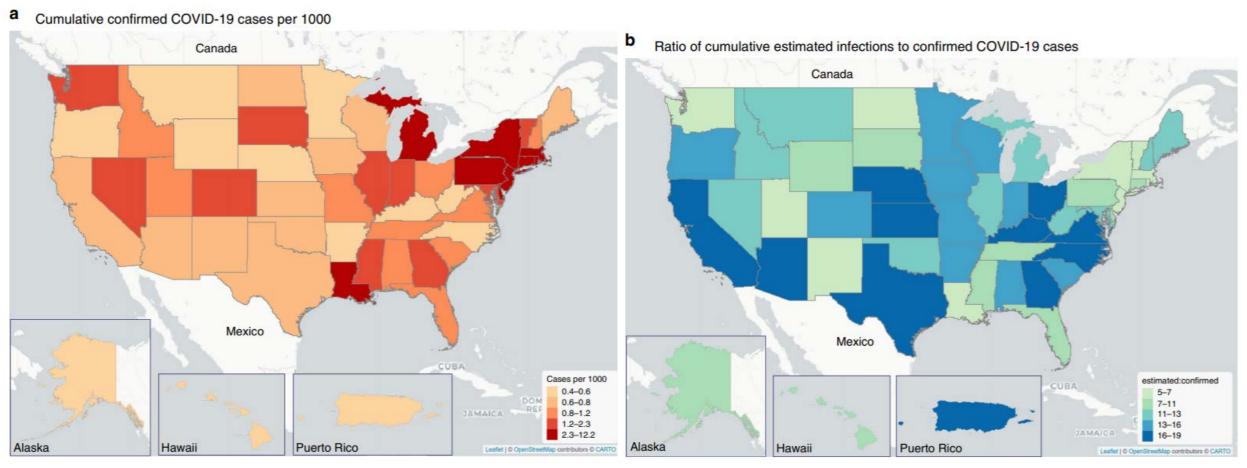
ARTICLE

https://doi.org/10.1038/s41467-020-18272-4

OPEN

Substantial underestimation of SARS-CoV-2

infection in the United States



### IS HOLIDAY TRAVEL SAFE?

### How long should adult working children Quarantine before they visit elderly parents over Christmas?

# COVID-19 Precautions – Holiday Safety



The safest way to celebrate the holidays this year is to celebrate with people in your household. If you do plan to spend the holidays with people outside your household, take steps to make your celebration safer.

#### Safest Options

Least Safe Options



Gather only with those in your household

Host a virtual celebration



Create a "holiday bubble" using social distancing and testing

 Bubble is only as strong as the weakest link



Gather with individuals outside your "bubble"

At a minimum, follow CDC recommendations

# COVID-19 Precautions – Spread of COVID-19 After a Gathering





# What are your recommendations for how to safely invite people into my home to do necessary repairs during the pandemic?

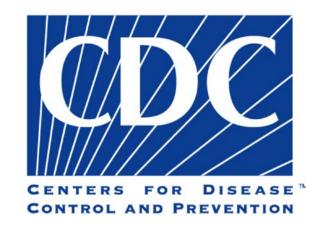
IN ADDITION TO OPENING WINDOWS, ETC. WHILE THE REPAIRS ARE ONGOING, SHOULD I DO ANY SPECIAL CLEANING AFTERWARD?

# COVID-19 Precautions – In-Home Services or Repairs



### What you need to know

- Ask the service provider to wear a <u>mask</u>.
- You and other household members should wear masks.
- Stay at least 6 feet from the service provider, and limit interactions between the service provider and other household members and pets.
- After the service is completed, <u>clean and disinfect</u> any surfaces in your home that may have been touched by the service provider.



# HOW LONG ARE YOU CONTAGIOUS TO OTHERS AFTER YOU TEST POSITIVE FOR COVID?

## COVID-19 Precautions -How Long Are You Contagious?



SARS-CoV-2, SARS-CoV, and MERS-CoV viral load dynamics. duration of viral shedding, and infectiousness: a systematic review and meta-analysis



Muge Cevik, Matthew Tate, Ollie Lloyd, Alberto Enrico Maraolo, Jenna Schafers, Antonia Ho

Background Viral load kinetics and duration of viral shedding are important determinants for disease transmission. Lance Microbe 2020 We aimed to characterise viral load dynamics, duration of viral RNA shedding, and viable virus shedding of severe Published Online acute respiratory syndrome coronavirus 2 (SARS-CoV-2) in various body fluids, and to compare SARS-CoV-2, November 19, 2020 SARS-CoV, and Middle East respiratory syndrome coronavirus (MERS-CoV) viral dynamics.

Methods In this systematic review and meta-analysis, we searched databases, including MEDLINE, Embase, Europe PubMed Central, medRxiv, and bioRxiv, and the grey literature, for research articles published between Jan 1, 2003, ofMedicins, Univenity of and June 6, 2020. We included case series (with five or more participants), cohort studies, and randomised controlled trials that reported SARS-CoV-2, SARS-CoV, or MERS-CoV infection, and reported viral load kinetics, duration of viral shedding, or viable virus. Two authors independently extracted data from published studies, or contacted authors to request data, and assessed study quality and risk of bias using the Joanna Briggs Institute Critical Appraisal Checklist Western General Hospital, tools. We calculated the mean duration of viral shedding and 95% CIs for every study included and applied the random-effects model to estimate a pooled effect stze. We used a weighted meta-regression with an unrestricted maximum likelihood model to assess the effect of potential moderators on the pooled effect size. This study is registered with PROSPERO, CRD42020181914.

Findings 79 studies (5340 individuals) on SARS-CoV-2, eight studies (1858 individuals) on SARS-CoV, and 11 studies (799 individuals) on MERS-CoV were included. Mean duration of SARS-CoV-2 RNA shedding was 17 · 0 days (95% CI 15-5-18-6; 43 studies, 3229 individuals) in upper respiratory tract, 14-6 days (9-3-20-0; seven studies, 260 individuals) in lower respiratory tract, 17-2 days (14-4-20-1; 13 studies, 586 individuals) in stool, and 16-6 days (3-6-29-7; two studies, 108 individuals) in serum samples. Maximum shedding duration was 83 days in the upper respiratory tract, 59 days in the lower respiratory tract, 126 days in stools, and 60 days in serum. Pooled mean SARS-CoV-2 shedding duration was positively associated with age (slope 0.304 [95% CI 0.115-0.493]; p=0.0016). No MRC-University of Glasgov study detected live virus beyond day 9 of illness, despite persistently high viral loads, which were inferred from cycle Contro for Virus Research, threshold values. SARS-CoV-2 viral load in the upper respiratory tract appeared to peak in the first week of illness, whereas that of SARS-CoV peaked at days 10-14 and that of MERS-CoV peaked at days 7-10.

interpretation Although SARS-CoV-2 RNA shedding in respiratory and stool samples can be prolonged, duration of viable virus is relatively short-lived. SARS-CoV-2 titres in the upper respiratory tract peak in the first week of illness. Early case finding and isolation, and public education on the spectrum of illness and period of infectiousness are key to the effective containment of SARS-CoV-2.

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determine the duration of infectiousness, which is a critical parameter to inform effective control measures detection has not been well characterised. A compre-(SARS-CoV-2) shedding, viral load dynamics and duration factors, such as age and disease severity, is lacking. piratory sampling, peak viral load was observed just RT-PCR, and viable virus shedding in various body fluids

before, or at the time of, symptom onset.24 Viral RNA Viral load kinetics and the duration of viral shedding are shedding was reported to be persistent in the upper important determinants for disease transmission. They respiratory tract and in faeces for more than 1 month after illness onset.1 However, the duration of SARS-CoV-2 RNA and disease modelling. Although several studies have hensive understanding of viral load dynamics, length of evaluated severe acute respiratory syndrome coronavirus 2 viral shedding, and how these measures relate to other

of viral shedding reported across studies so far have been We aimed to characterise the viral load dynamics heterogeneous.1 In several case series with serial res- of SARS-CoV-2, duration of viral RNA shedding by

ttps://doi.org/10.1016

(M Covik MRCP); NHS Lothia Edinburgh, UK (M Covik, opiratory Medicine, Oueen Edinburgh Medical School College of Medicine and Diseases, Cotuono Hospital A ORN dei Colli, Naples, Italy (A EMaraolo MD); and

Dr Muge Cevik, Division of Infection and Clobal Health Research, School of Medicine University of St Andrews Fife KY16 9TE UK mc349@st-andrews.ac.uk

	Classification of illness severity	Median (IQR*) duration of SARS-CoV-2 positivity in cohort, days	Viral dynamics in patients with severe illness vs those with non-severe illness	p value
Chen et al (2020)⁵	ICU vs non-ICU patients	11 (95% CI 10-12)	Median time to viral clearance significantly longer in ICU vs non-ICU patients (HR 3·17, 95% CI 2·29–4·37)	Only HR provided
Chen et al (2020) <sup>26</sup>	China CDC guideline (version 7)	12 (8–16)	Shedding duration varies by severity: asymptomatic 6 days; mild 10 days; moderate 12 days; serious 14 days; critical 32 days	<0.0001
Tan et al (2020) <sup>17</sup>	China CDC guideline (version 6)	Nasopharyngeal swab: 12 (range 3–38); any sample: 22 (range 3–38)	Viral shedding significantly longer in patients with severe illness: any sample 23 days vs 20 days (note that nasopharyngeal swab 14 vs 11 days was non-significant)	0.023 (any sample)
Xu et al (2020) <sup>36</sup>	WHO criteria	17 (13-32)	Higher proportion of patients with severe illness had shedding > 15 days (34·2% vs 16·2%)	0.049
Yan et al (2020) <sup>37</sup>	China CDC guideline (version 6)	23 (18–32)	No difference in shedding duration (general illness 23 days vs severe illness 26 days vs critical illness 28 days)	0.51
Zheng et al (2020) <sup>23</sup>	China CDC guideline (version 6)	Respiratory sample: 18 (13-29)	Shedding duration significantly longer in patients with severe illness (21 vs 14 days) in respiratory samples; no difference in shedding duration in stool or serum samples	0.04

CDC=Center for Disease Control and Prevention. HR=hazard ratio. ICU=intensive care unit. SARS-CoV-2=severe acute respiratory syndrome coronavirus 2. \*IQR unless otherwise stated.

Table 1: Severity of illness and viral dynamics

"Although SARS-CoV-2 RNA shedding in respiratoryand stool samples can be prolonged, duration of viable virus is relatively short-lived. SARS-CoV-2 titres in the upper respiratory tract peak in the first week of illness."

www.thelancet.com/microbe Published online November 19, 2020 https://doi.org/10.1016/52666-5247(20)30172-5

# DO YOU AGREE WITH THE UPDATED CDC QUARANTINE GUIDELINES?

# COVID-19 Precautions – Quarantine Guidelines

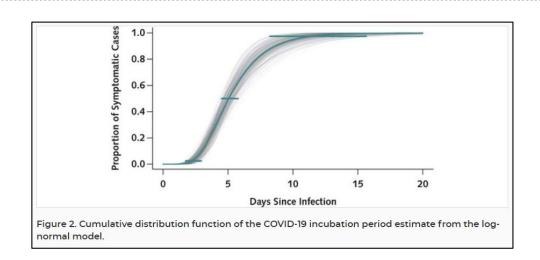


The efficacy of quarantine to reduce the transmission of infectious disease depends on how compliant people are. Is a shorter quarantine time with higher compliance more effective than the "gold standard" with lower compliance?

Length of Quarantine	Estimated Post- Quarantine Residual Transmission Risk
14 days	0-3%
10 days	1-10%
7 days w/ negative test	5-12%

"The current recommended 14-day quarantine period is the most conservative regimen and should be maintained whenever possible." – Greater Houston Area Public Health Authority

"If we can increase compliance with quarantine but also make sure we have a quarantine policy that allows us to catch as many cases as we can, that's the best of both worlds." - Assn. Public Health Officials



ANY NEW RECOMMENDATIONS FOR SCHOOLS? ANY UPDATES TO TRANSMISSION OF THE VIRUS AMONG TEENS? IS CONTACT TRACING AMONG TEENS FRUITFUL? ARE QUARANTINE TIMES APPROPRIATE?

What is the impact of COVID on toddlers – risk of sending young children to small school settings? What is the likelihood of children transferring COVID to elder family members?

# AAMC Report: Kids, Schools and COVID-19: What we know – and what we don't



- The virus spreads in schools but schools are rarely superspreaders
- School outbreaks typically come from the community not vice versa
  - Many school outbreaks have been tied to social gatherings among school children outside of school
- Children transmit the virus but not as often as adults do
  - Children carry large amounts of the virus in their respiratory systems,
     they transmit the virus, but not as effectively as adults
- Behavioral challenges matter sometimes as much as biology
  - Mask compliance, washing hands and maintaining distance



ACTIVELY ENGAGED IN PHYSICAL THERAPY AT METHODIST ORTHOPEDIC, NOTHING URGENT, SHOULD I STOP GOING TO THAT AND ANYTHING INVOLVING A GYM UNTIL THE SPRING OF NEXT YEAR?

# ACTIVELY ENGAGED IN PHYSICAL THERAPY AT METHODIST ORTHOPEDIC, NOTHING URGENT, SHOULD I STOP GOING TO THAT AND ANYTHING INVOLVING A GYM UNTIL THE SPRING OF NEXT YEAR?

It is necessary to continue to receive any medical treatments, preventive and therapeutic. Houston Methodist is safe and you should continue to access medical care when you need it!



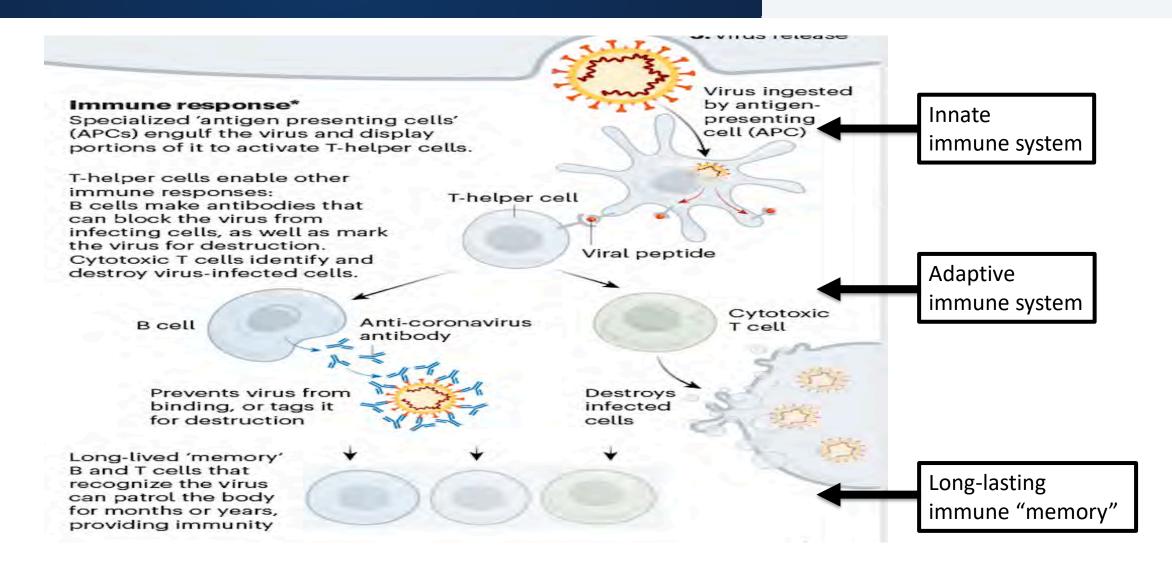
# COVID-19 Vaccines – Update Town Hall December 10, 2020

H. Dirk Sostman, MD FACR



### How Immunity Develops





# Vaccine Progress: Frontrunners



Vaccine	Antibody Response (Phase 1 - 2)	T Cell Response (Phase 1 – 2)	Technology	N of Doses	EUA Target
Moderna	100% (2x – 8x CP)	100%	mRNA	2	December 17, 2020
Pfizer / BioNTech	100% (5x – 30x CP)	94%	mRNA	2	December 10, 2020 Granted in UK December 1 & in Canada December 9

CP = convalescent plasma

# Vaccine Progress: Second Wave



Vaccine	Antibody Response	T Cell Response	N of Doses	Technology	EUA Target
Oxford / Astra Zeneca	100% (= CP)	100%	2	Adenovirus Ad5 vector	December 2020 (UK) Q1 2021 (US)
Novavax	100% (2x CP)	100% (subgroup)	2	Peptide / nanoparticle	Q1 2021
181	100%	82%	1	Adenovirus Ad26 vector	Q1 2021
Inovio	94% overall	94% overall	2	DNA + electroporation	Q1 2021?
COVAXX				Synthetic peptide multi-epitope	Still in Phase 1
COVI-Vac				Intranasal live attenuated vaccine	Starting Phase 1
And many others, e.g.:	Sputnik V ICL	CanSino Sanofi, GSK	CNBG1 CNBG2	Novartis Medicago	Curevac Sinovac

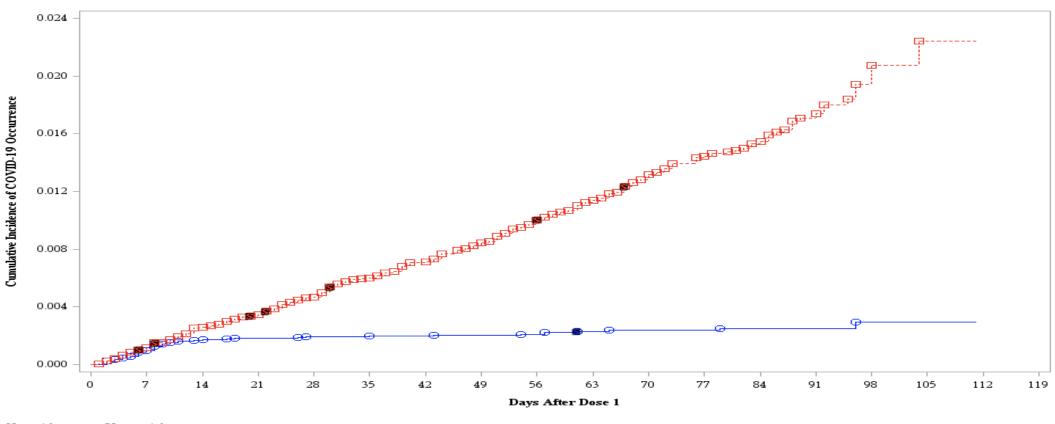
https://www.nytimes.com/interactive/2020/science/coronavirus-vaccine-tracker



#### Pfizer

- 43,538 enrolled and randomized to vaccine or placebo
- 170 infections so far, 162 in placebo group → 95% protection
- 10 severe cases, 9 in placebo group → 90% protection
- Appears to be effective in all age and ethnic groups
- Unknowns
  - Prevents infection or only symptomatic illness? (key issue for transmission)
  - Duration of protection?
- Safety profile appears excellent
  - Reactogenic effects
  - Two allergic reactions in UK?
- Logistics: Two doses & extreme cold chain (stable for 6 mos at -70° C, 5 days at 2-8° C, 2 hrs at 25° C)





No. with events/No. at risk

A: 0/21314 21/21230 37/21054 39/20481 41/19314 42/18377 42/17702 43/17186 44/15464 47/14038 48/12169 48/9591 49/6403 49/3374 50/1463 50/398 50/0 B: 0/21258 25/21170 55/20970 73/20366 97/19209 123/18218 143/17578 166/17025 192/15290 212/13876 235/11994 249/9471 257/6294 267/3301 274/1449 275/398 275/0

— O A: BNT162b2 (30 μg) ---- B: Placebo

Note: "S" indicates subjects with severe COVID-19 or COVID-19 leading to hospitalization.

PFIZER CONFIDENTIAL SDTM Creation: 17NOV2020 (10:49) Source Data: adc19ef Table Generation: 17NOV2020 (21:40)

(Cutoff Date: 14NOV2020, Snapshot Date: 16NOV2020) Output File: ./nda2\_unblinded/C4591001\_Efficacy\_FA\_164/adc19ef\_f001\_km\_d1\_aai



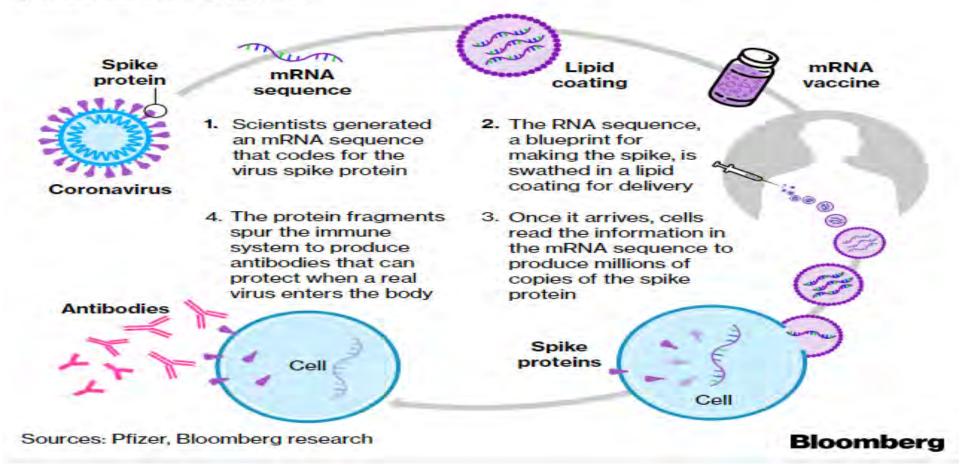
#### Moderna

- > 30,000 enrolled and randomized to vaccine or placebo
- 196 infections so far, 185 in placebo group → 94.1% protection
- 30 severe cases, all in placebo group → 100% protection
- Similar efficacy in all groups (preliminary review)
- Unknowns
  - Prevents infection or only symptomatic illness? (key issue for transmission)
  - Duration of protection?
- Safety profile appears excellent (few details yet)
- Logistics: Two doses & cold chain needed (stable in freezer for 6 mos, refrigerator for 30 days, room temp for 12 hrs)



#### **How mRNA Vaccines Work**

The vaccine spurs healthy cells to produce viral proteins that stimulate a potent immune response



#### Vaccine Updates: Pfizer

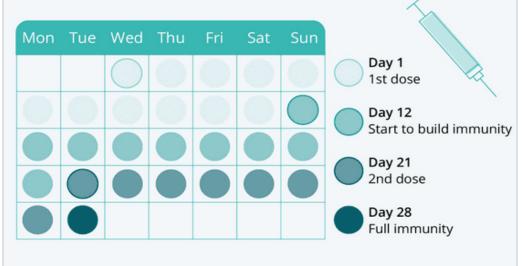


# The Pfizer/BioNTech Vaccination Process

Posted on DailyShot.com 04-Dec-2020

@SoberLook

Vaccination process for the Pfizer/BioNTech Covid-19 BNT162b2 vaccine



Source: Pfizer/BioNTech via BBC









Margaret Keenan, 91 yo, first clinical Pfizer vaccination

# Adverse Reactions: Severe "Reactogenic" Effects



- More likely after second dose
- Controllable with anti-inflammatories (Tylenol, Motrin, etc.)
- High fever (39 40° C) in 2%
- Pfizer (company press release November 18, 2020)
  - Fatigue (3.8%)
  - Headache (2%)
- Moderna (company press release November 16, 2020)
  - Fatigue (9.7%)
  - Muscle pain (8.9%)
  - Joint pain (5.2%)
  - Headache (4.5%)

# Pfizer Vaccine FDA Safety Overview



Adverse Effect (AE)	Vaccine Group	Placebo Group
Solicited inject site AE	73%	11%
Solicited systemic AE	70%	34%
Unsolicited non-serious AE	27%	13%
Serious AE	0.6%	0.5%
Withdrawal for AE	0.6%	0.5%
Allergic reaction	0.6%	0.5%
Death	2	4

# Pfizer Vaccine 18 – 55 yo Group Mild – Moderate "Reactogenic" Effects



Adverse Effect	Vaccine Group	Placebo Group
Fever < 39° C	14%	0.3%
Fatigue	55%	22%
Headache	49%	23%
Chills	33%	4%
GI	12%	10%
Myalgia	35%	8%
Arthralgia	21%	5%
Pain / Anti-inflammatory Medication	45%	13%

# Pfizer Vaccine > 55 yo Group Mild - Moderate "Reactogenic" Effects

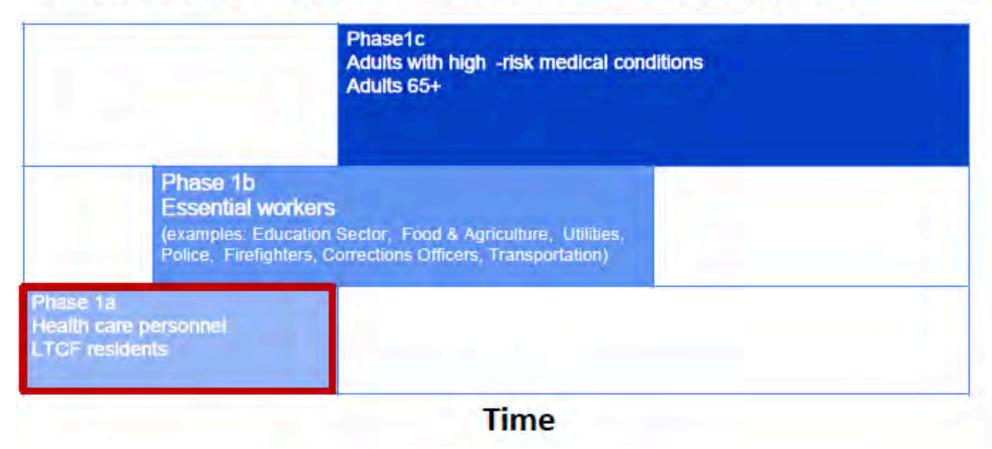


Adverse Effect	Vaccine Group	Placebo Group
Fever < 39° C	11%	0.2%
Fatigue	48%	17%
Headache	38%	14%
Chills	22%	3%
GI	9%	6%
Myalgia	28%	5%
Arthralgia	18%	4%
Pain / Anti-inflammatory Medication	38%	10%

#### **ACIP** Report to CDC



## Work Group Proposed Interim Phase 1 Sequence



#### **UK Government**



#### The <u>first phase</u> of the programme

#### Offer vaccination

- 1 Residents in a care home for older adults and their carers
- All those 80 years of age and over. Frontline health and social care workers
- 3 All those 75 years of age and over
- All those 70 years of age and over. Clinically extremely vulnerable individuals
- 5 All those 65 years of age and over
- All individuals aged 16 years to 64 years with underlying health conditions which put them at higher risk of serious disease and mortality
- 7 All those 60 years of age and over
- 8 All those 55 years of age and over
- 9 All those 50 years of age and over





- **Protecting health care workers** who fill a critical role in caring for and preserving the lives of COVID-19 patients and maintaining the health care infrastructure for all who need it.
- Protecting frontline workers who are at greater risk of contracting COVID-19 due to the nature of their
  work providing critical services and preserving the economy.
- **Protecting vulnerable populations** who are at greater risk of severe disease and death if they contract COVID-19.
- Mitigating health inequities due to factors such as demographics, poverty, insurance status and geography.
- **Data-driven allocations** using the best available scientific evidence and epidemiology at the time, allowing for flexibility for local conditions.
- **Geographic diversity** through a balanced approach that considers access in urban and rural communities and in affected ZIP codes.
- Transparency through sharing allocations with the public and seeking public feedback.

#### **Houston Methodist**



- Vaccine Scientific Committee advice on priority tiers
  - Tier 1:
    - Health care workers with high-risk exposure (3 sub-tiers)
    - Government mandates (e.g., Long Term Care, first responders)
  - Tier 2:
    - Health care workers with moderate-risk exposure
  - Tier 3:
    - Health care workers with low-risk exposure and general population
      - HCWs likely will be prioritized for reasons of operational efficiency
  - Within Tiers, if vaccine is limited, prioritize based on health risks (age & co-existing conditions)

#### Obstacles?



- Reluctance to accept vaccination
  - Political issues
  - Concerns about potential side effects
- Logistics challenges
  - Supplies (borosilicate glass vials, needles, syringes, etc.)
  - Cold chain of refrigeration
  - Air freight capacity (8,000 jumbo jets)
  - Paperwork, customs, health regulations, etc.
  - Organizing administration sites, records, personnel
  - Monitoring safety, side effects

#### When Will We Get Back to Normal?



- If we want things to remain the same, some things will have to change:
  - Masks and preventive measures needed until population is immune and transmission controlled
  - Even then, masks in flu season or high-risk situations will still be advisable
- If all goes well, population immunity is likely by end of Q2 or Q3:
  - A big "IF"
  - Schools return in the fall
  - Travel and leisure gradually return to something resembling antebellum "normal"
- Learning from our mistakes we must:
  - Rebuild early warning / early response capabilities
  - Invest in vaccine and infectious disease research
  - Repair communication among government, health care and public
  - Seek dialogue and consensus about health priorities and social contracts
  - Foster trust in science and understanding of people's needs and work



# THANK YOU FOR ATTENDING OUR TOWN HALL CONVERSATION

If you would like more information about the Houston Methodist DeBakey Heart & Vascular Center, the Department of Surgery or The Society for Leading Medicine, please contact foundation@houstonmethodist.org

Take care and be well

