# HHP/HPH COVID-19 Community Webinar Series

Thursday, June 17, 2021 5:30pm – 6:30pm





# Moderator - 06/17/21

Andy Lee, MD

Medical Director, Hawai'i Health Partners
Chief of Staff, Pali Momi Medical Center
Hawai'i Pacific Health



### Disclaimer:

 The following is intended as information resource only for HHP/HPH providers, clinicians, administrative and clinical leaders.

 Specific areas may not pertain directly to your clinical practice area and/or may not be applicable to your practice based on your existing workflows, infrastructure, software (e.g. EHR), and communications processes.

# Webinar Information

- You have been automatically muted.
   You cannot unmute yourself.
- You will be able to submit questions via the Q&A section.
  - Due to time constraints, any unanswered questions will be addressed this week and posted on the HHP website
- A recording of the meeting will be available tomorrow on the HHP website and intranet.



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### 1. Step 1: Confirm your attendance

 You should have completed a brief questionnaire before joining today's live webinar.

### 2. Step 2: HPH CME team will email you instructions

- Complete and submit evaluation survey that will be emailed to you within one week of the offering.
- Your CE certificate will be immediately available to you upon completion of your evaluation.
- Questions? Email <u>hphcontinuingeduc@hawaiipacifichealth.org</u>



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### **Disclosures**

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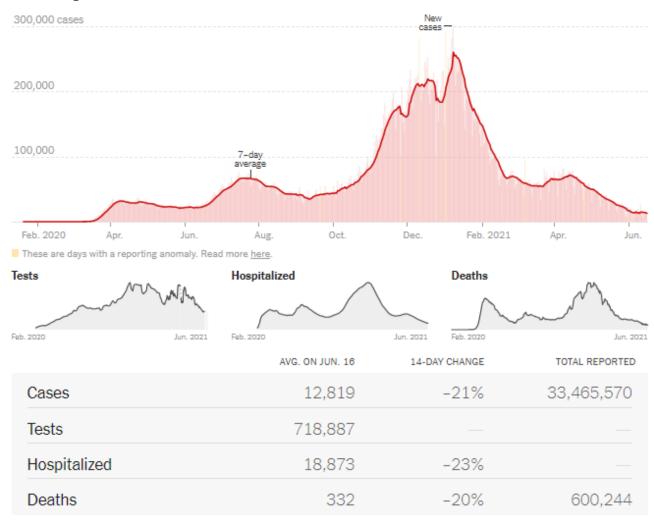
# COVID-19 Updates

Gerard Livaudais, MD, MPH
Executive Vice President, Population
Health and Provider Networks,
Hawai'i Pacific Health



# **United States**

### New reported cases

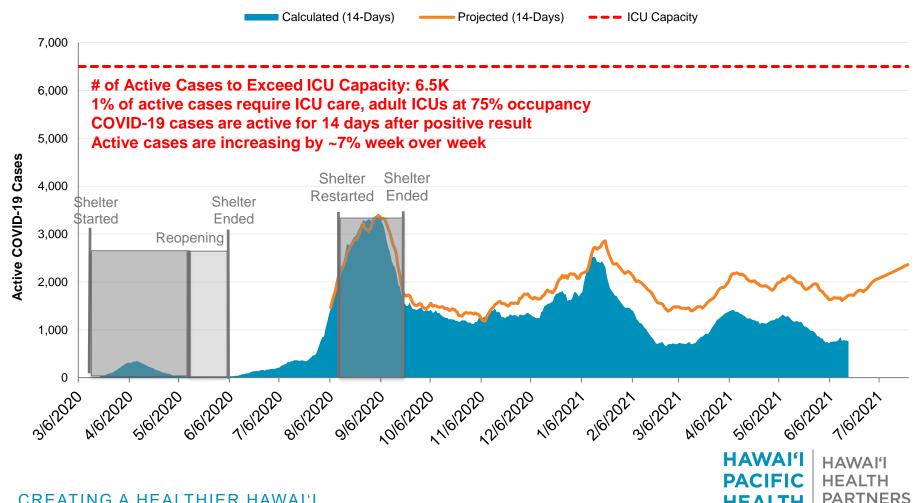


https://www.nytimes.com/interactive/2021/us/covid-cases.htm



# Projected Active COVID-19 Cases

#### Hawaii Actual v. Projected Active COVID Cases Updated 6/17/2021



**HEALTH** 

As of 6/17/21	Total Census	ICU beds occupied	# Ventilators in use	# New Admissions w/ positive COVID-19	# Patients currently hospitalized w/ suspect or confirmed COVID-19	# Patients currently on a ventilator w/ confirmed COVID-19	# Patients currently in ICU w/ confirmed COVID-19
KMCWC	161	69	18	0	S: 0 C: 0	0	0
РММС	106	14	5	1	S: 0 C: 4	1	1
SMC	134	9	7	0	S: 0 C: 1	0	0
WMC	64	2	0	0	S: 0 C: 0	0	0

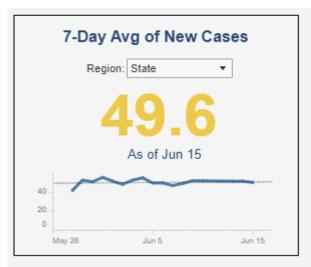
S = Suspected; C= Confirmed

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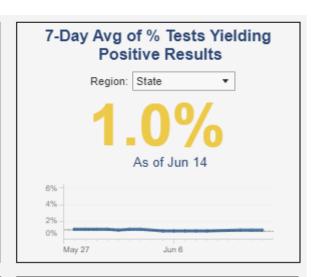
		Setting and severity of illness				
		Ambulatory care: mild-to- moderate disease	Hospitalized: mild-to-moderate disease without need for suppl. oxygen	Hospitalized: severe but non- critical disease (SpO <sub>2</sub> ≤94% on room air)	Hospitalized: critical disease (e.g., in ICU needing MV, or septic shock, ECMO)	
1	Hydroxy- chloroquine (HCQ)*	NA	Recommend against use  ⊕⊕⊕○	Recommend against use  ⊕⊕⊕○	Recommend against use  ⊕⊕⊕○	
2	HCQ*+ azithromycin	NA	Recommend against use	Recommend against use	Recommend against use	
3	Lopinavir + ritonavir	NA	Recommend against use	Recommend against use  ⊕⊕⊕○	Recommend against use  ⊕⊕⊕○	
4-6	Corticosteroids	NA	Suggest against use ⊕○○○	Suggest use  The state of the s	Recommend use  Commend use  R: If dexamethasone is unavailable, equivalent total daily doses of alternative glucocorticoids may be used.**	
7	Tocilizumab	NA	NA	Suggest use  ⊕⊕⊖⊖  R: Patients, particularly those who response to steroids alone, who put a high value on avoiding possible adverse events of tocilizumab and a low value on the uncertain mortality reduction, would reasonably decline tocilizumab.  R: In the largest trial on the treatment of tocilizumab, criterion for systemic inflammation was defines as CRP ≥75 mg/L	Suggest use  ⊕⊕⊖⊖  R: Patients, particularly those who response to steroids alone, who put a high value on avoiding possible adverse events of tocilizumab and a low value on the uncertain mortality reduction, would reasonably decline tocilizumab.  R: In the largest trial on the treatment of tocilizumab, criterion for systemic inflammation was defines as CRP ≥75 mg/L	
8-9	Convalescent plasma	Recommended only in the context of a clinical trial (knowledge gap)	Suggest against use  ⊕⊕○○	Suggest against use ⊕⊕○○	Suggest against use ⊕⊕○○	
10- 12	Remdesivir		Suggest against routine use ⊕○○○	Suggest use ⊕⊕⊕⊖	Routine initiation of remdesivir:	
		NA		5 days vs. 10 days, on supplemental oxygen but without mechanical ventilation or ECMO: Suggest use	Suggest against use ⊕○○○	

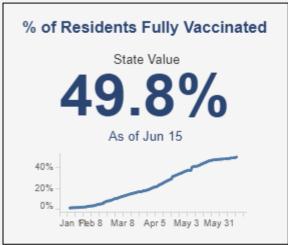
13	Famotidine	NA	Suggests against use except in a clinical trial	Suggests against use except in a clinical trial	Suggests against use except in a clinical trial
14	Bamlanivimab + etesevimab or casirivimab + imdevimab	Suggest use  R: Patients with mild to moderate COVID-19 who are at high risk of progression to severe disease admitted to the hospital for reasons other than COVID-19 may also receive bamlanivimab/etesevimab or casirivimab/imdevimab. Local variant susceptibility may be considered in the choice of the most appropriate neutralizing antibody therapy. There are limited data on efficacy of bamlanivimab/etesevimab or casirivimab/imdevimab in high- risk patients between 12 and 18 years of age.	NA	NA	NA
15	Bamlanivimab monotherapy	NA	NA	Recommend against use ⊕⊕⊕⊖	NA
16	Baricitinib + Remdesivir	NA	NA	Suggest use  R: Baricitinib 4 mg per day up to 14 days or until discharge from hospital.  R: Baricitinib appears to demonstrate the most benefit in those with severe COVID-19 on high-flow oxygen/non- invasive ventilation at baseline.	
17	Baricitinib + remdesivir + corticosteroids	NA	NA	Suggest use****  R: Baricitinib 4 mg daily dose for 14 days or until hospital discharge. The benefits of baricitinib plus remdesivir for persons on mechanical ventilation are uncertain.	NA
18- 19	Ivermectin	Suggests against use except in a clinical trial	NA	Suggests against use except in a clinical trial ⊕○○○	NA

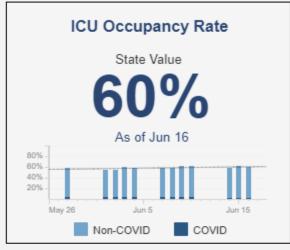
# **COVID Pau Dashboard**

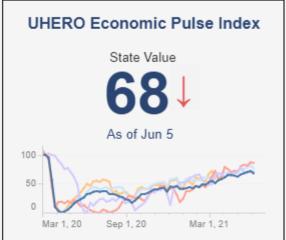








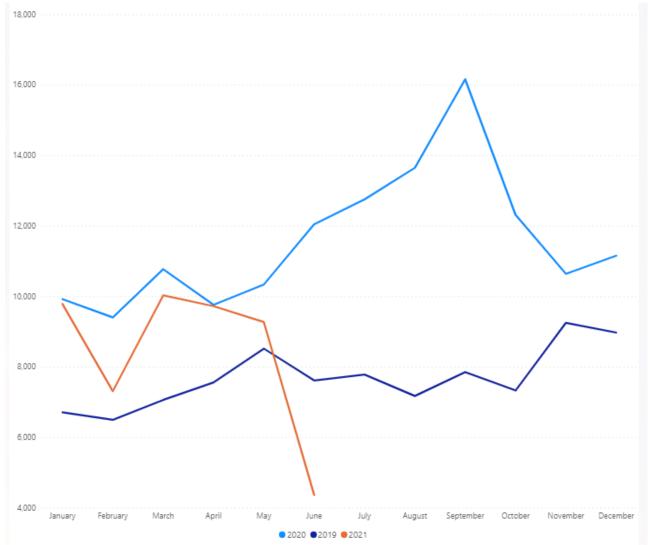




#### UHERO.data



### All Hawai'i CARES Call Volumes by Month, 2019-2021





### Hawaii COVID-19 Vaccine Summary

LAST UPDATED ON Wednesday, June 16, 2021

#### All persons age 12 and older are eligible for vaccination

#### **DOSES ADMINISTERED**

STATEWIDE TOTAL VACCINES ADMINISTERED

1,621,282

TOTAL STATEWIDE PROGRESS

55% completed 61% initiated

FEDERAL PHARMACY PROGRAM

283,244

FEDERAL AGENCY DOSES

161,883

JURISDICTION DOSES\*

1,176,155

#### STATE PROGRESS

Status Completed Population Total Population



Fully Vaccinated
Individuals
(State Benchmarks)

55% of Total Population



785,746 Completed Doses

Click to view Dose Tracking

#### **NAVIGATE TO OTHER VIEWS**

Click buttons to navigate to other views



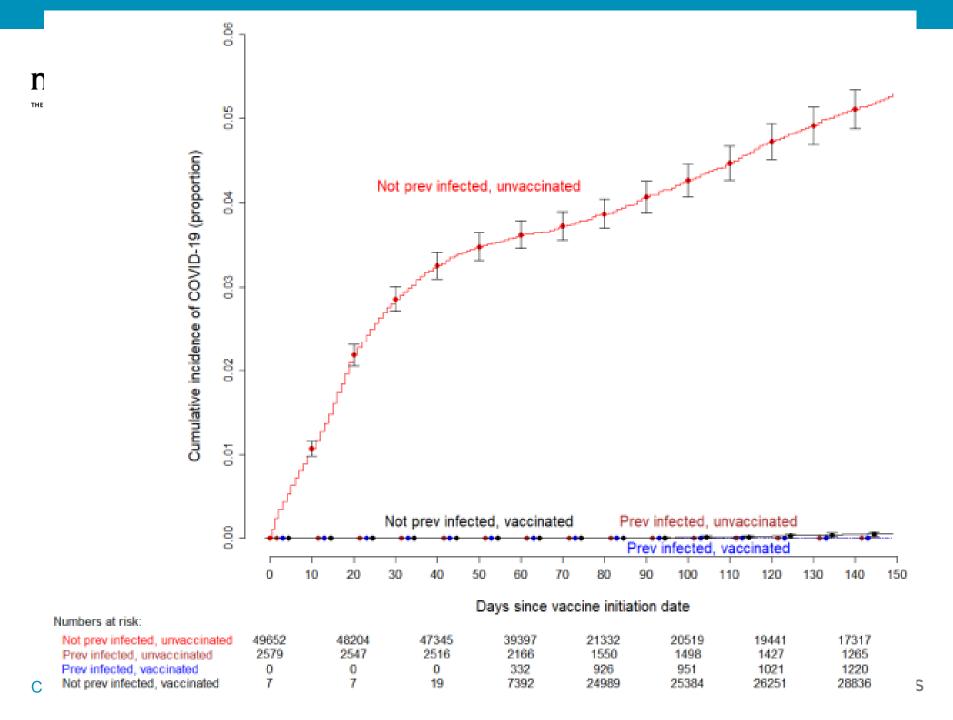
DOSE TRACKING

COUNTY

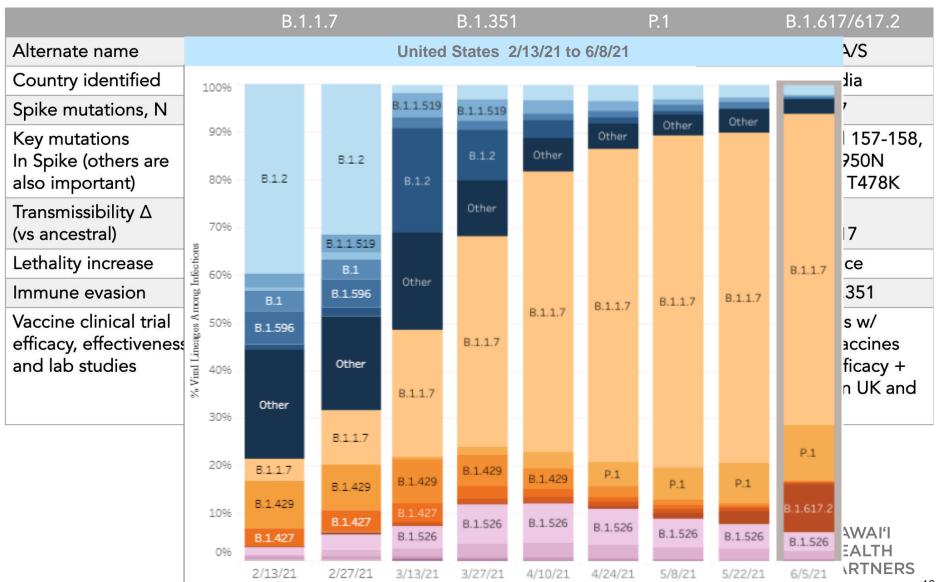
MAP

AGE

RACE

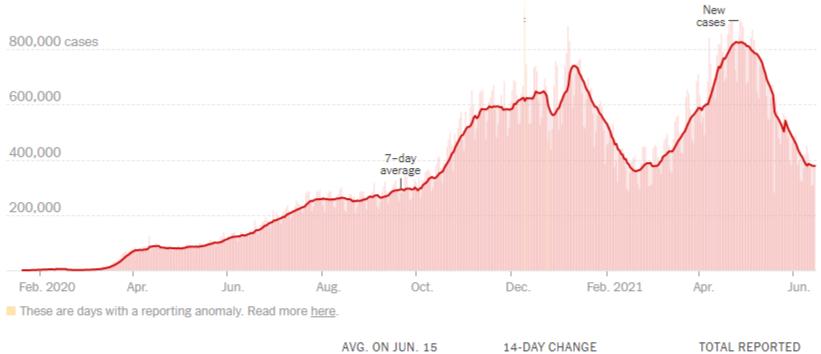


# Variants of Concern



# Worldwide

### New reported cases



	AVG. ON JUN. 15	14-DAY CHANGE	TOTAL REPORTED
Cases	378,028	-21%	176,630,053
Deaths	10,818	-6%	3,822,553

About this data

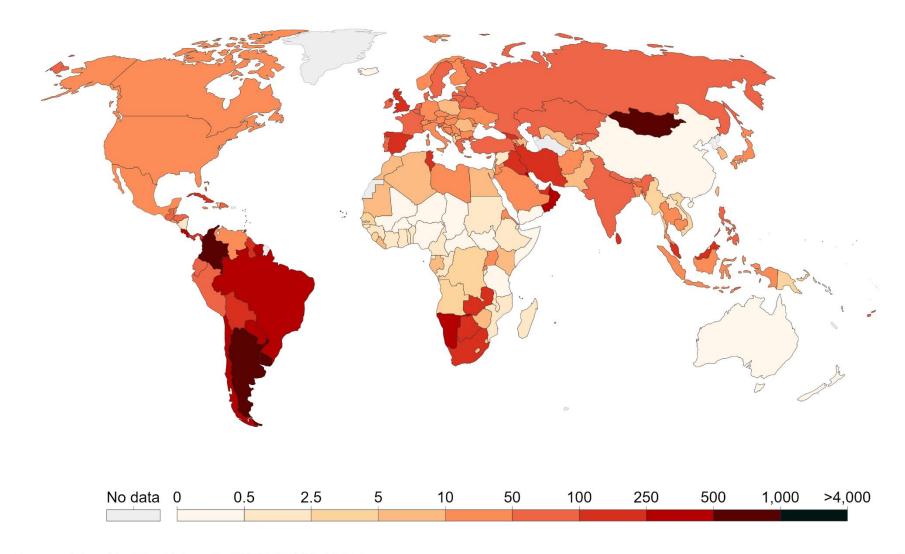
https://www.nytimes.com/interactive/2020/world/coronavirus-maps.html

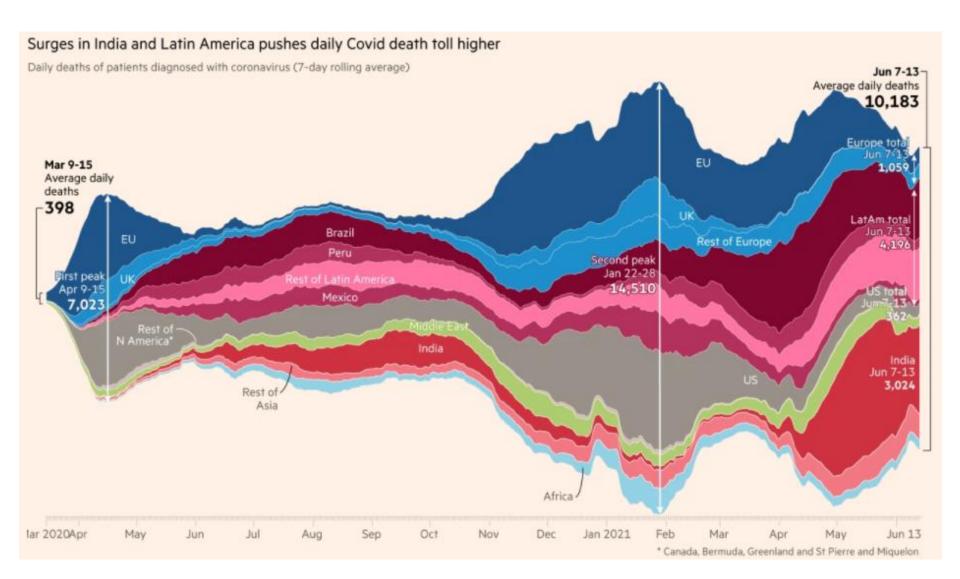
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PACIFIC HEALTH
HEALTH PARTNERS

### Daily new confirmed COVID-19 cases per million people



Shown is the rolling 7-day average. The number of confirmed cases is lower than the number of actual cases; the main reason for that is limited testing.





https://www.ft.com/content/a2901ce8-5eb7-4633-b89c-cbdf5b386938



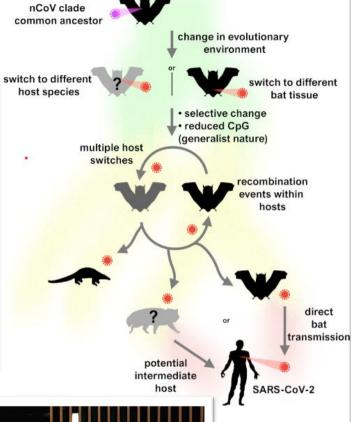
# **COVID-19 Origin Theories**



# Biden Orders Intelligence Inquiry Into Origins of Virus

The directive came as health officials and scientists have renewed calls for a more rigorous examination.

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https://www.youtube.com/watch?v=IdYDL\_RK--w

# Y.C. James "Jimmy" Yen



Volunteered in WW1 for the Y.M.C.A. among 20,000 illiterate Chinese laborers who had been imported to dig trenches. While writing letters home for them by day and translating news for them at night, he developed a basic Chinese vocabulary of about 1,300 characters.

#### China:

- Trained illiterate farmers to record births and deaths, vaccinate against smallpox and other diseases, give first aid and health education talks, and help communities keep their wells clean.
  - These services were delivered Barefoot Doctors in communities where the infant mortality was more than 200 deaths per 1,000 live births and life expectancy was only 35 years. Estimated 90% reduction in endemic Schistosomiasis.

#### World:

 Barefoot Doctor concept gained attention around the world, served as a guiding concept for CHW programs in many countries, including Honduras, India, Indonesia, Tanzania, and Venezuela.



# Updates on Long COVID and the HPH Long COVID Care Program



Bennett Loui, MD

Primary Care Provider – Internal Medicine, Straub Medical Center, Hawai'i Pacific Health Chief of Internal Medicine, Hawai'i Pacific Health Medical Group



# Long COVID

### Background

- Post-COVID conditions are associated with a spectrum of physical, social, and psychological consequences, as well as functional limitations
- Impact on societal health and economy is extensive
- Knowledge about these conditions is emerging

### Definitions

- Acute COVID-19: symptom onset to 2-4 weeks
- Long COVID: 4+ weeks after onset
  - Long Hauler
  - Post-Acute COVID19 Syndrome
  - Post COVID Conditions (CDC)
  - Post Acute Sequalae of SARS-CoV2 (NIH PASC Initiative)



Detection unlikely PCR positive PCR negative **Fatigue** Joint pain Dyspnea Nasopharyngeal Cough Anxiety/depression Viral isolation from respiratory tract **PTSD** Headaches **Palpitations** Chest pain SARS-CoV-2 Thromboembolism exposure Hair loss Week -2 Week -1 Week 1 Week 2 Week 3 Week 4 Week 12

Decline in quality of life Muscular weakness

Persistent oxygen requirement

Sleep disturbances Cognitive disturbances (brain fog)

Chronic kidney disease

6 months

Before symptom onset After symptom onset

Nalbandian, A., Sehgal, K., Gupta, A. et al. Post-acute COVID-19 syndrome. Nat Med 27, 601-615 (2021). https://doi.org/10.1038/s41591-021-01283-z



Viral load

MEDICAL GROUP

# Long COVID Epidemiology

- Prevalence reports vary due to differing methods
  - which symptoms, how reported, which patients
- Michigan
  - 60 days post discharge following hospital treatment of COVID: 32.6% of patients reported persistent symptoms
- Wuhan
  - 6 months post discharge: majority still reported 1+ symptoms
- UK COVID Symptom Study App
  - 28 days post diagnosis: 13% had symptoms
  - 8 weeks: 4.5% had symptoms
- Fairhealth claims review (6/15/2021)
  - 1,959,982 COVID patients, 30+ days after onset/diagnosis:
    - 23% sought evaluation for conditions relatable to LC
    - Asymptomatic COVID patients: 19% sought evaluation
  - Limitation- not compared to non-COVID patients
- 1. Chopra etal Ann Intern Med https://doi.org/10.7326/m20-5661 (2020)
- 2. Huang etal Lancet 397, 220-232 (2021)
- 3. https://www.fairhealth.org/



# Long COVID Epidemiology

- High risk for Long COVID
  - Severe acute illness (ICU)
  - High BMI
  - Older age
  - Women

- History of respiratory disease
- History of anxiety or depression
- Proposed pathophysiologic mechanisms
  - Persistent viral reservoir
  - Inflammatory injury
  - Microvascular injury, hypercoagulability
  - Immune system dysregulation
  - Post-intensive care syndrome (PICS)



# Long COVID Symptoms and Conditions by System

#### General

Symptoms of Fatigue, post-exertional malaise (PEM), fever

### Pulmonary

- Symptoms of difficulty breathing, cough
- Hypoxemia, fibrosis
- PFTs and imaging may not show abnormalities

#### Cardiovascular

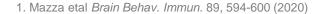
- Symptoms of palpitations, chest pain
- Stress cardiomyopathy (echocardiogram)
- Myocarditis (cardiac MRI)
- Dysautonomia, Catecholaminergic state (Holter/Zio)
- Abstinence from athletics

### Neurological

- Symptoms of brain fog, dizziness, headache, anosmia
- Hypoxic-ischemic injury, microvascular thrombosis, viral CNS infection
- Brain imaging often normal

### Psychiatric

Depression or anxiety (25-56%), PTSD, insomnia





# Long COVID Symptoms and Conditions by System

- Hematologic
  - PE, VTE <5%, prophylaxis under investigation</li>
- Dermatologic
  - Hair loss (20%)
- Rheumatologic
  - Reactive arthritis
- Renal
  - New CKD (13%)
- Endocrine
  - DKA, thyroiditis/hypothyroidism
- MIS-C
  - Multisystem Inflammatory Syndrome in Children
  - May present 4 weeks or more after COVID infection
- Social/Occupational
  - Loss of ability to work
  - Financial stress



# Long COVID Management Themes

- Laboratory and diagnostic testing abnormalities may be absent
- Consider conservative diagnostic approach in the first 4-12 weeks
- Patients need reassurance and validation
- Set achievable goals, focus on specific symptoms and functions, engage with support groups
- Current understanding is incomplete and guidance will change



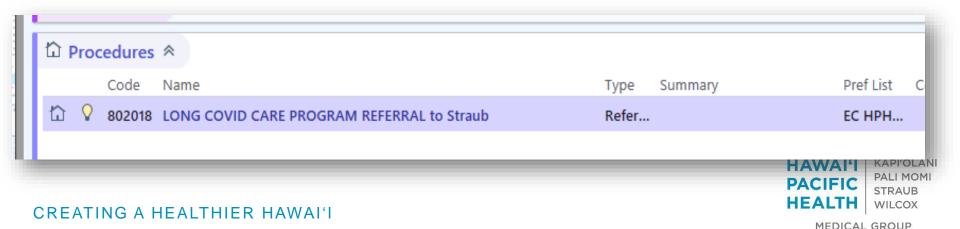
# Long COVID Guidance and Support

- Studies worldwide and reviews are numerous
- CDC Post-COVID Conditions Interim Guidance
  - Posted this week and presented today 6/17/2021
    - https://www.cdc.gov/coronavirus/2019-ncov/hcp/clinical-care/post-covid-index.html
  - Provides a framework for healthcare providers in their initial assessment, evaluation, management, and followup of persons with possible post-COVID conditions
- Patient advocacy
  - COVID Advocacy Exchange
  - Facebook groups
  - Survivor Corps



# Long COVID Care Program (LCCP)

- Goal is to support the care of patients experiencing persistent symptoms after the acute phase of COVID-19
- Referrals
  - Handoff from HHP Covid Virtual Monitoring Program (now closed)
  - Referral from a HHP PCP or Specialist providers
  - Now open to referrals from community providers
  - Epic order: LONG COVID CARE PROGRAM REFERRAL to Straub
  - Fax form available at HPH website



ONG COVID CAF	re program i	REFERRAL to Straub					✓ Accept	A Carr
Referral:	To provider:		00					
	To dept:	TELEHEALTH - SC 🔎	, ,					
	Reason:	Continuity of Care	Specialty Services Requ	red Continuit	ty of Care			
	Priority:		specialty services requ	Continui	ty or care			
	Type:	Routine P	Committee of Committee of					
	.,,,	Consult/Test/Trea	Consult/Test/Treat					
Reason for Refer	rral							
Active symptoms	s or			J				
Date of COVID-1 diagnosis ©	19	Ė						
Is patient on oxy	/gen Yes	No						
Pertinent special	lty providers on	care team, if applicable						
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Comments:	Assc	Encounter Diagnoses	C	des Qualifier	Comment			
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# HAWAI'I PACIFIC HEALTH LONG COVID CARE PROGRAM REFERRAL

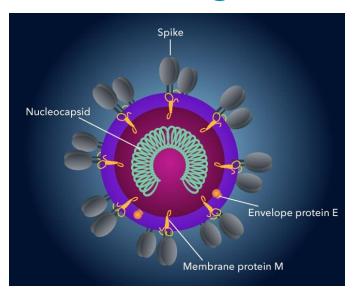
Fax: 808-957-0685 | Phone: 808-462-5459

rom:	Date:
Phone: Fax:	No. of Pages (including this page):
Referring PCP:	
PATIENT INFORMATION:	
Patient Name:	
DOB: Pho	one:
Address:	
nsurance Provider:	Member ID
Referral/Authorization Attached? ☐ Yes ☐ No	
Date of COVID-19 Diagnosis:	
Reason for Referral / Lasting Symptoms:	
□ On O2 □ Shortness of Breath	
☐ Snortness of Breath ☐ Fatique or Tiredness	
☐ Joint Pain	
□ Chest Pain	
Cough	
<ul> <li>□ Difficulty with Thinking and Concentration</li> <li>□ Depression</li> </ul>	
☐ Muscle Pain	
□ Headache	
☐ Intermittent Fever	
☐ Fast-beating or pounding heart, or palpitations	
Other	



# Referrals to Long COVID Care Program

 Positive COVID-19 test not required but clinical suspicion should be high



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- Consider Antibody test
  - SARS-CoV2 IgG (anti-N protein) does NOT turn positive after mRNA/J+J vaccinations
  - Usually develops by 14 days after onset of symptoms
  - Duration of detectable antibody uncertain
  - Rare cross-reactivity with other coronaviruses has been reported

# Long COVID Care Program Process

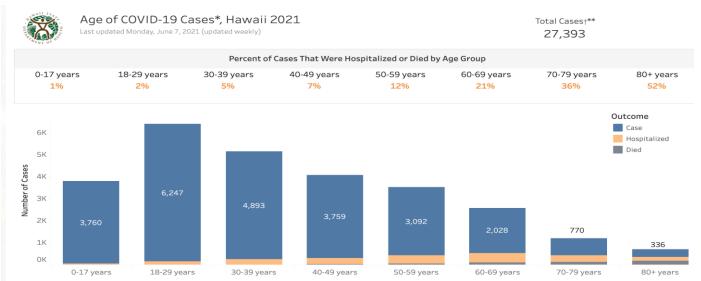
- Evaluation and Monitoring by Telephone or Video visits
  - LCCP NP coordinates closely with PCP
  - Intensity of monitoring individualized based on clinical needs
- Interdisciplinary Collaboration
  - Leaders in key medical specialties have been engaged
    - Cardiology, Pulmonology, Neurology, Psychology, Infectious Disease
    - Advise management, create protocols, facilitate referrals
    - Pediatric patients are evaluated by a general Pediatrician with specialist assistance as needed
  - Coordination with Case Management, PT/OT and Complex Care Team



### **LCCP Patients Referred**

54 patients, 51 from HPHMG

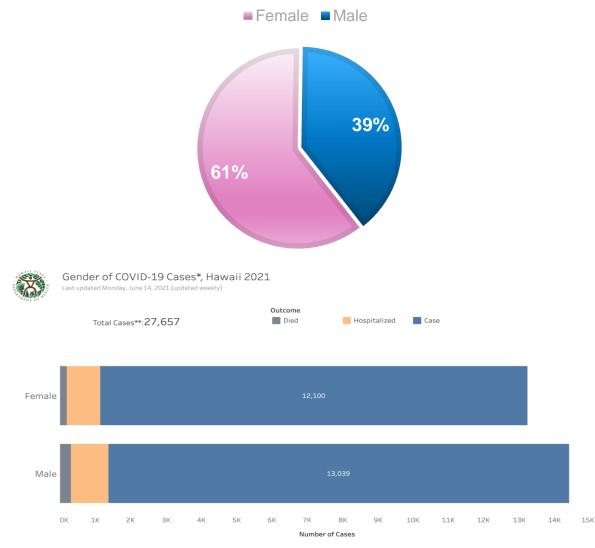








# **LCCP Patients Referred**





### LCCP Experience: Most Commonly Addressed Conditions



### **LCCP** Management

- Conservative diagnostic approach in weeks 4-12
  - Labs
  - Imaging
  - Other evaluation
    - EKG, Echocardiogram, Holter/Zio, PFT
- Home O2 weaning
- Breathing exercises (Stasis, Johns Hopkins)
- Advocacy group resources
- Discuss vaccinations
  - COVID-19, Pneumovax
- Referrals
  - PT
  - Social work
  - Specialist providers



### LCCP Outcomes

- Follow up intensity
  - Monitoring at intervals typically between 1 and 6 weeks
  - Most patients continue to be followed
  - Some have been discharged for follow up with PCP
- A database of LC patients is being maintained for future outreach
- Feedback from providers and patients have been very positive



### LCCP: The Future

- COVID-19 case rates decreasing and LCCP referral rates are decreasing
- Many may not be familiar with LC or the LCCP
  - 37K Hawai'i cases, 34M USA cases, 170M Worldwide to date
- We are promoting awareness internally and to community
  - List of COVID patients by PCP distributed to providers March 2021
  - Website, TV, Print media, Webinars
- LCCP will continue to update knowledge and resources to support providers



# Long COVID: The Future

- How to prevent Post COVID? Vaccinate.
- How to treat Long COVID? Maybe vaccinate too.
  - Specific treatments may not yet exist, but the effect of vaccination is being studied

WILCOX

MEDICAL GROUP

- 40% of respondents to an internet poll reported symptom improvement <a href="https://www.survivorcorps.com/yale">https://www.survivorcorps.com/yale</a>
- Yale has launched a formal study
   <a href="https://www.yalemedicine.org/news/vaccines-long-covid">https://www.yalemedicine.org/news/vaccines-long-covid</a>
- Active and future research on Long COVID is critical to improving understanding and guiding clinical practice

# Thank you!

#### HPH Long COVID Care Program Team

Nohea Taufaasau NP Sheri Yoshino NP Mary Yoshida RN Bennett Loui MD

Sam Evans MD
Huidy Shu MD
Bart Pillen PhD
Sanah Christopher MD

Heidi Hillesland MD Michaella Okihara MD Geldilyn Ebbay RN BSN



# Myocarditis Associated with the COVID-19 Vaccine



#### Andras Bratincsak, MD, PhD

Pediatric and Adult Congenital Cardiologist, Hawai'i Pacific Health Medical Group

Assistant Professor of Pediatrics, University of Hawai'i, John A. Burns School of Medicine, Department of Pediatrics



#### FDA NEWS RELEASE

# Coronavirus (COVID-19) Update: FDA Authorizes Pfizer-BioNTech COVID-19 Vaccine for Emergency Use in Adolescents in Another Important Action in Fight Against Pandemic



For Immediate Release: May 10, 2021

Español

Today, the U.S. Food and Drug Administration expanded the emergency use authorization (EUA) for the Pfizer-BioNTech COVID-19 Vaccine for the prevention of coronavirus disease 2019 (COVID-19) caused by severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) to include adolescents 12 through 15 years of age. The FDA amended the EUA originally issued on Dec. 11, 2020 for administration in individuals 16 years of age and older.



### **COVID-19 Vaccine Statistics**

		<b>US</b> (as of 6/3/21)	<b>Hawaii</b> (as of 6/1/21)	<b>HPH</b> (as of 6/2/21)
12-17 years old	At least 1 dose	6,393,198*	33,470	
	2 doses	2,328,858*	10,029	
12-18 years old	At least 1 dose			10,142
	2 doses			3,400

<sup>\*</sup>Number of vaccines administered in the US is a presumed number calculated based on the following information for number of vaccines administered in population aged 12 years and older and 18 years and older presented on the CDC website.

- ≥ 12 years old: 168,954,018 received at least 1 dose, 136,635,500 received 2 doses
- $\geq$  18 years old: 162,560,820 received at least 1 dose, 134,306,642 received 2 doses



Advanced Search

#### Vaccines & Immunizations



Clinical Considerations: Myocarditis and Pericarditis after Receipt of mRNA COVID-19 Vaccines Among Adolescents and Young Adults

#### Summary

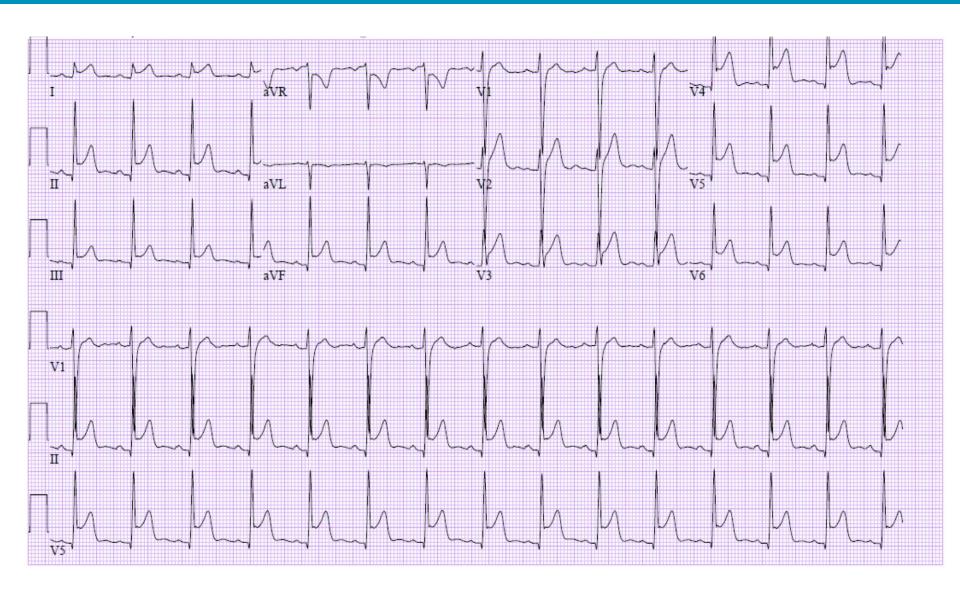
Since April 2021, increased cases of myocarditis and pericarditis have been reported in the United States after mRNA COVID-19 vaccination (Pfizer-BioNTech and Moderna), particularly in adolescents and young adults. There has not been a similar reporting pattern observed after receipt of the Janssen COVID-19 Vaccine (Johnson & Johnson).



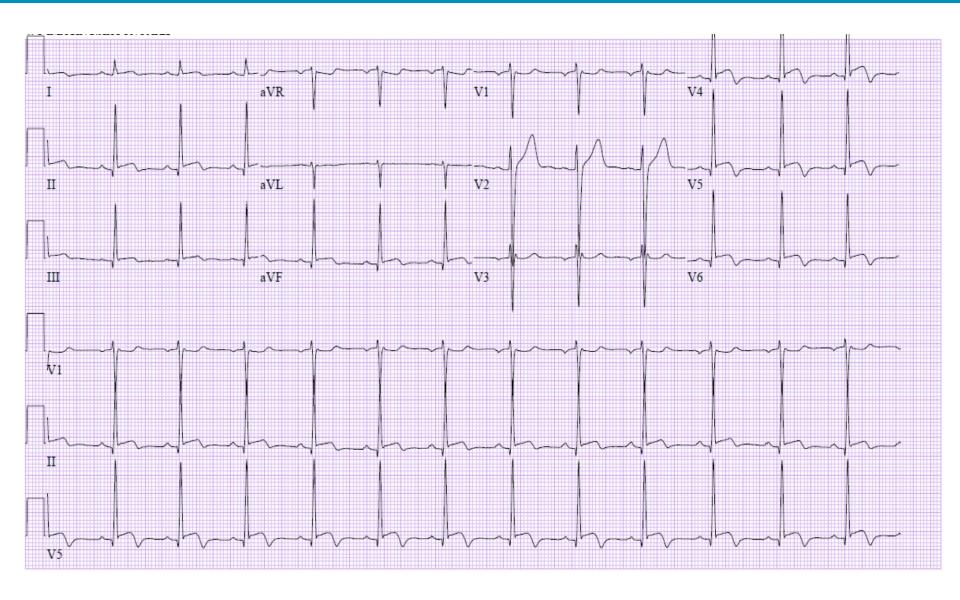
# 3 Cases with Myocarditis Within 3 Weeks

- 15 yo M presenting with acute chest pain
  - -3 days following 1st dose of COVID-19 mRNA vaccine (Pfizer)
  - Had headache, dry cough, tactile fever the day after vaccination.
  - Troponin T: 304 ng/L; CKMB: 54.6 U/L
- 16 yo M with acute chest pain
  - -2 days following 2nd dose of COVID-19 mRNA vaccine
  - No significant symptoms immediately following vaccination
  - Troponin T: 431 ng/L; CKMB: 39.3 U/L
- 17 yo M with acute chest pain
  - -2 days following 2nd dose of COVID-19 mRNA vaccine
  - Had headache, pain over inoculation site, mild fever the day after vaccination
  - Troponin T: 749 ng/L; CKMB: 46.5 U/L

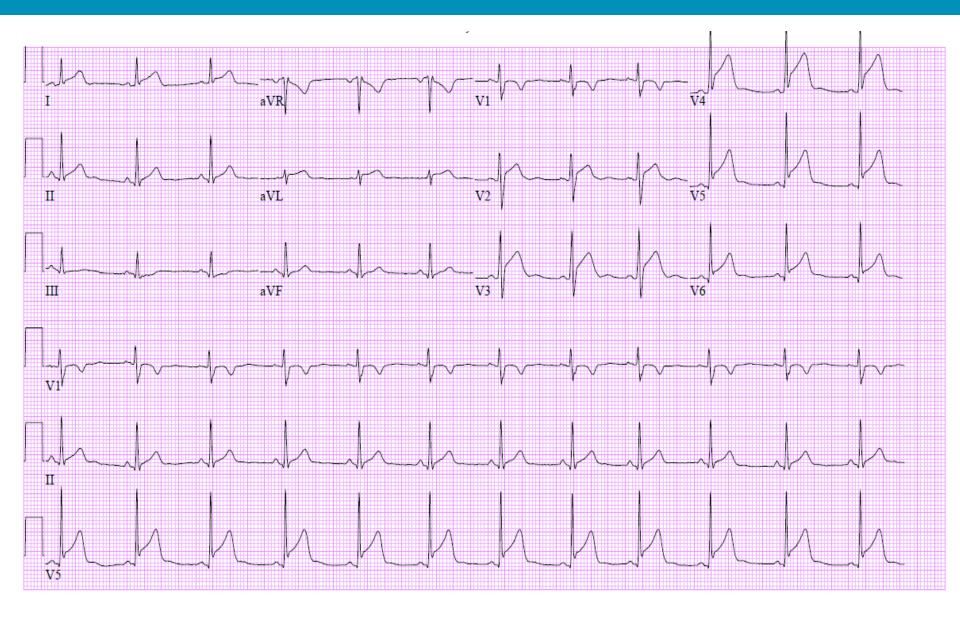












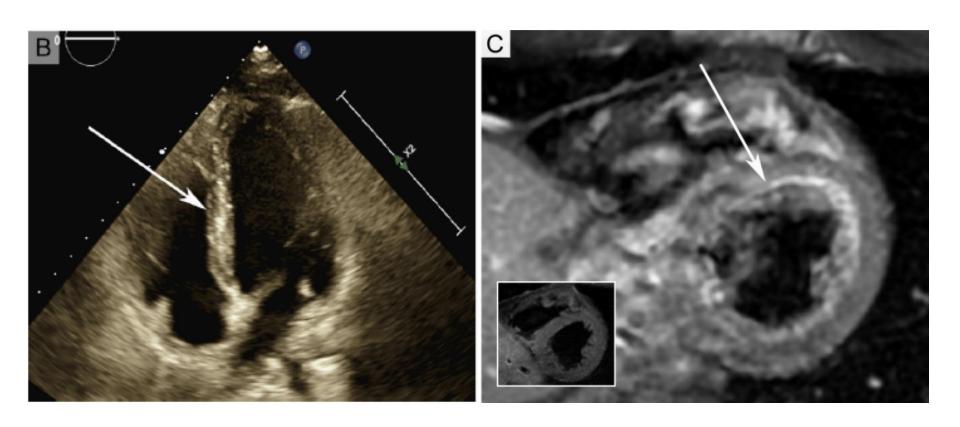


			Patient A	Patient B	Patient C
Demographics	Age, years		15	16	17
	Sex		Male	Male	Male
	Ethnicity/Race		Filipino	Filipino, Caucasian	Hawaiian/Pacific Islande
History of cardiac cor	nditions		None	None	None
BNT162b2 vaccine	Number of doses given		1	2	2
	Days after last vaccine admir pain	nistration to onset of chest	3	2	2
Vital signs at	Heart rate, beats per minute		116	108	78
presentation	Blood pressure, mmHg		119/72	131/91	120/71
	Temperature, °C		37.2	37.2	36.6
	Respiratory rate		18	20	20
	SpO2 in room air, %		96	100	98
Cardiac markers	Troponin T, ng/L (<20 ng/L)	At presentation	304	431	749
(normal range)		Peak	832	1210	938
	CKMB, U/L (<4.1 U/L)	At presentation	54.6	39.3	46.5
		Peak	54.6	39.3	46.5
	NT-proBNP, pg/mL (0-125	At presentation	108	325	55
	pg/mL)	Peak	498	364	111
Other laboratory	ESR, mm/hr (0-15 mm/hr)		7	31	N/A
results (normal	Procalcitonin (<0.10 ng/mL)		0.15	<0.06	N/A
range)	CRP, mg/L (<5.0 mg/L)		18.5	24.3	N/A
	SARS-CoV-2 RT-PCR		Negative	Negative	Negative
	SARS-CoV-2 IgG (<1.40 inde	ex)	Negative (0.24)	Negative (0.03)	Negative (0.02)
	Respiratory viral panel PCR		Negative	Negative	Negative



	Patient A	Patient B	Patient C
Length of stay, days	3	2	2
Electrocardiogram findings	ST segment elevation in lateral leads, left axis deviation	ST segment elevation in inferolateral leads, T wave inversion	ST segment elevation in lateral leads
Echocardiogram findings	Increased echogenicity of myocardium, LVEF 63%	LVEF 60%	LVEF 63%
Cardiac MRI findings	Not obtained	Global early gadolinium enhancement ratio of 4.0 (positive >4.0), no significant T2 hyperintense myocardial edema, no delayed myocardial enhancement suggestive of myocardial necrosis	Not obtained







# Accepted for Publication

Self-limited myocarditis presenting with chest pain and ST segment elevation in adolescents after vaccination with the BNT162b2 mRNA vaccine

Journal:

Cardiology in the Young

Jihyun Park, Dona Brekke, Andras Bratincsak



### Report on myocarditis following the Pfizer vaccine

# PEDIATRICS

OFFICIAL JOURNAL OF THE AMERICAN ACADEMY OF PEDIATRICS

# Symptomatic Acute Myocarditis in Seven Adolescents Following Pfizer-BioNTech COVID-19 Vaccination

Mayme Marshall, MD, Ian D. Ferguson, MD, Paul Lewis, MD, MPH, Preeti Jaggi, MD, Christina Gagliardo, MD, James Steward Collins, MD, Robin Shaughnessy, MD, Rachel Caron, BA, Cristina Fuss, MD, Kathleen Jo E. Corbin, MD, MHS, Leonard Emuren, MBBS, PhD, Erin Faherty, MD, E. Kevin Hall, MD, Cecilia Di Pentima, MD, MPH, Matthew E. Oste, MD, MPH, Elijah Paintsil, MD, Saira Siddiqui, MD, Donna M. Timchak, MD, Judith A. Guzman-Cottrill, DO



Table 1. Demographic and clinical characteristics of seven cases of symptomatic myocarditis after dose #2 of Pfizer-BioNTech COVID-19 vaccine

	Patient 1	Patient 2	Patient 3	Patient 4	Patient 5	Patient 6	Patient 7
Age (years)	16	19	17	18	17	16	14
Sex	Male	Male	Male	Male	Male	Male	Male
Race/Ethnicity	White	White	White	White	Latino	White	White
Weight (kg)	68	68	71	69	64	71	92
BMI (kg/m²)	24	19	21	21	19	22	28
Exposure to	None	None	None	None	None	None	None
COVID-19 in 14 days prior to illness onset							
Time between vaccine dose	2	3	2	2	4	3	2
#2 and symptom onset (days)							
Total hospital LOS (days)	6	2	2	4	5	3	4
ICU LOS (days)	4	None	None	4	5	2	2
		Svr	nptoms Upon F	Presentation			
Chest pain	Present	Present	Present	Present	Present	Present	Present
Other pain	Bilateral arm	Myalgias	Bilateral		Bilateral arm		
•	pain	, ,	arm pain, numbness, paresthesia		pain, abdominal pain		
Fever	38.3° C by history	Subjective, chills		Subjective	Subjective		38.3° C by history
Fatigue	Present	Present		Present			
Other	Nausea, vomiting, anorexia,	Weakness		Nausea	Nausea, vomiting, anorexia, SOB,	SOB	SOB

<sup>--:</sup> Not present; Kg: kilograms, BMI: Body Mass Index, LOS: Length of Stay; ICU: Intensive Care Unit; SOB: Shortness of breath.



Table 2. Summary of diagnostics and therapeutics: seven cases of symptomatic myocarditis after dose #2 of Pfizer-BioNTech COVID-19 vaccine

	Patient 1	Patient 2	Patient 3	Patient 4	Patient 5	Patient 6	Patient 7
		Lah	oratory Findings o	n Admission			
Troponin (ng/mL) (normal range)	Troponin I: 2.59 (<0.03)	High-sensitivity troponin T: 232 (< 14)	Troponin I: 5.55 (<0.045)	Troponin T: 1.09 (<0.01)	Troponin T: 3.2 (<0.01)	Troponin T: 0.66 (<0.01)	Troponin I: 22.1 (<0.045)
Brain natriuretic peptide (pg/mL) (normal < 100)							107.9
NT pro-BNP (pg/mL) (normal < 125)	428		376		978	149	
Peripheral white blood cell count (thousand/cu mm)	6.97	8.69	11.8	12.6	16.3	5.0	8.11
Absolute lymphocyte count (thousand/cumm)	1.69	1.39	2.13	2.3	4.1	1.4	1.05
Absolute neutrophil count (thousand/cu mm)	4.65	5.93	7.46	9.5	9.8	2.8	4.73
Platelet count (thousand/cu mm)	198	208	231	236	297	189	208
Albumin (g/dL)	3.9	4.1	4.1	4.4	4.0	3.8	3.5
Aspartate transaminase (units/L)	54	29	41	82	150	59	87
Alanine transaminase (units/L)	30	14	33	20	46	22	38
Ferritin (ug/L)	70		90	103	347	65	84



C-reactive protein	0.99	6.7	2.5	12.7	18.1	1.5	7.7
(mg/dL)							
(normal < 1.0)							
Erythrocyte	18	13	6	40	38	3	10
sedimentation rate							
(mm/hr)							
Prothrombin time			14.0		12.1	11.4	14.8
(seconds)							
Partial thromboplastin	22.3		31.4		30.4	27.9	35.6
time (seconds)							
International	1.11		1.06		1.13	1.06	1.2
Normalized Ratio INR							
		Othe	er Pertinent Laborato	ry Findings			
Highest troponin	Troponin I:	High sensitivity	Troponin I:	Troponin T:	Troponin T:	Troponin T:	Troponin I:
(ng/mL)	12.43 (<0.80)	Troponin T: 388	12.20 (<0.045)	1.09 (<0.01)	3.33 (<0.01)	0.82 (<0.01)	22.1 (<0.045)
(normal range)		(<14)					
Lowest troponin prior	Troponin I: 1.42		Troponin I:	Troponin T: 0.4	Troponin T:	Troponin T:	Troponin I:
to discharge (ng/mL)	(<0.80)		5.79 (<0.045)	(<0.01)	0.96 (<0.01)	0.01 (<0.01)	8.02 (<0.045)
(normal range)							
Highest BNP							205 pcg/mL
(normal range)							(<100)
Highest NT-pro BNP	482 pg/mL		376 pg/mL (<300)		978 pcg/mL	275 pcg/mL	
(normal range)	(<125)				(<125)	(<125)	
Highest C-reactive	1.23	6.7	2.53	12.7	18.1	1.8	12.7
protein (mg/dL)							
(normal < 1.0)							
COVID-19 PCR	Negative	Negative	Negative	Negative	Negative	Negative	Negative
COVID-19 spike			Positive	Positive	Positive	Positive	
antibody			(Roche)	(Roche)	(Roche)	(Roche)	
(Manufacturer)							
COVID-19 nucleocapsid	Negative		Negative	Negative	Negative	Negative	Negative
antibody	(Abbott)		(Roche)	(Roche)	(Roche)	(Roche)	(Abbott)
(Manufacturer)							



	†		1	1		†	1
Respiratory pathogen	Negative	Negative	Negative	Negative	Negative	Negative	Negative
panel PCR*	(BioFire)	(BioFire)	(BioFire)	(BioFire)	(BioFire)	(BioFire)	(BioFire)
(Manufacturer)							
Adenovirus diagnostics	Negative serum		Negative serology	Negative serum	Negative		Negative serum
	PCR			PCR	serum PCR		PCR
Enterovirus diagnostics	Negative serum		Negative serology	Negative serum	Negative	Negative	Negative serum
	PCR			PCR	serum PCR	serum PCR	PCR
Cytomegalovirus	Negative serum		Negative serology	Negative serum	Negative	Negative	Negative
diagnostics	PCR			PCR	serum PCR	serum PCR	serology
Epstein-Barr virus			Negative serology	Negative serum	Negative	Negative IgM,	Negative
diagnostics				PCR	serum PCR	positive IgG	serology
						antibody	
Other diagnostics			Negative		Negative	Negative Lyme	Negative
			Parvovirus,		Parvovirus	serology,	Parvovirus IgM,
			Bartonella, and		and Bartonella	negative	positive
			Lyme serology,		serology,	Mycoplasma	Parvovirus IgG
			negative urine		negative HHV-	serum PCR,	antibody,
			drug screen		6 serum PCR	negative	negative
						Parvovirus	Mycoplasma PCR
						serum PCR	(throat swab)
			l Diagnostic Imaging Fi	ndings		Jeruii i ek	(till out swab)
Cardiac MRI	LGE		LGF		LGE	LGE, diffuse	LGE
Cardiac IVIKI		LGE involving mid		Fibrosis,			
	(subepicardial)	LV wall,	(subepicardial)	myocardial	(epicardial)	myocardial	(subepicardial)
	involving lateral	myocardial edema	involving basal	edema,	involving	edema	involving mid and
	LV apex,	of basal	anterolateral and	hyperemia, mild	anterior and		apical LV free
	myocardial edema	inferolateral LV	basal to mid-	mitral	lateral LV wall,		wall, myocardial
	of lateral LV wall,	wall	ventricular	regurgitation	no myocardial		edema,
	left axillary		inferolateral LV	(RF ~18%)	edema		hyperemia
	adenopathy		segments,				
			myocardial				
			edema, elevated				
			extracellular				
			volume fraction				
			(29.2%)				



Echocardiogram	Normal	Normal	Borderline basal lateral and basal posterior strain	Normal	Normal	Normal	Mildly depressed RV and LV systolic function (LVEF 47%)
Electrocardiogram	Atrioventricular dissociation with junctional escape rhythm, ST elevation	ST segment elevation (diffuse)	ST elevation (diffuse), T wave abnormality Therapeutics	ST elevation	Sinus bradycardia, T wave abnormality	ST elevation (diffuse)	ST elevation, low voltage of extremity leads
	T.,					T	LENG
Oxygen supplementation	None	None	None	None	None	None	LFNC
Vasoactive medications or inotropic support	None	None	None	None	None	None	None
Anti-inflammatory agents and other relevant medications	NSAID, IVIG, IV methylprednisolo ne, PO prednisone, famotidine	NSAID, colchicine, aspirin	NSAID, famotidine	NSAID, IVIG, IV methylprednisol one, PO prednisone	NSAID, IVIG, IV methylpreniso lone, PO prednisone, aspirin	IVIG, PO prednisone	NSAID, famotidine, furosemide

--: Not done; LGE: late gadolinium enhancement; LV: Left ventricular; RV: Right ventricular, LVEF: Left ventricular ejection fraction; LFNC: Low flow nasal cannula; NSAID: Non-steroidal anti-inflammatory drug; IVIG: Intravenous immunoglobulin; IV: intravenous; PO: per os (oral); q12hr: every 12 hours; HHV-6: Human herpesvirus-6



	Vaccine related myocarditis
Fever	no
Rash	no
Cough	no
Headache	maybe
Chest pain	YES
GI symptoms	no
Symptoms within a few days from mRNA vaccine	YES
Symptoms within a few days to COVID exposure	no
Symptoms within a few weeks to COVID exposure	no



	Vaccine related myocarditis	Acute COVID infection
Fever	no	maybe
Rash	no	maybe
Cough	no	YES
Headache	maybe	maybe
Chest pain	YES	no
GI symptoms	no	no
Symptoms within a few days from mRNA vaccine	YES	no
Symptoms within a few days to COVID exposure	no	YES
Symptoms within a few weeks to COVID exposure	no	no



	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
Fever	no	maybe	YES
Rash	no	maybe	YES
Cough	no	YES	maybe
Headache	maybe	maybe	YES
Chest pain	YES	no	no
GI symptoms	no	no	YES
Symptoms within a few days from mRNA vaccine	YES	no	no
Symptoms within a few days to COVID exposure	no	YES	no
Symptoms within a few weeks to COVID exposure	no	no	YES



	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
Fever	no	maybe	YES
Rash	no	maybe	YES
Cough	no	YES	maybe
Headache	maybe	maybe	YES
Chest pain	YES	no	no
GI symptoms	no	no	YES
Symptoms within a few days from mRNA vaccine	YES	no	no
Symptoms within a few days to COVID exposure	no	YES	no
Symptoms within a few weeks to COVID exposure	no	no	YES



# Laboratory Work-up and Comparison

	Vaccine related myocarditis
ESR / CRP elevation	YES
CK-MB	YES
Troponin	YES
SARS-CoV-2 PCR	no
SARS-CoV-2 IgG	no
Respiratory viral panel	no



# Laboratory Work-up and Comparison

	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
ESR / CRP elevation	YES	YES	YES
CK-MB	YES	no	maybe
Troponin	YES	no	YES (20-60%)
SARS-CoV-2 PCR	no	YES	maybe (60%)
SARS-CoV-2 IgG	no	maybe	YES
Respiratory viral panel	no	YES	no



# Laboratory Work-up and Comparison

	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
ESR / CRP elevation	YES	YES	YES
CK-MB	YES	no	maybe
Troponin	YES	no	YES (20-60%)
SARS-CoV-2 PCR	no	YES	maybe (60%)
SARS-CoV-2 IgG	no	maybe	YES
Respiratory viral panel	no	YES	no



# **Imaging Studies**

	Vaccine related myocarditis
ST segment elevation	YES
Decreased LV function	no
Coronary dilation	no
Cardiac MRI gadolinium enhancement	YES
Cardiac catheterization	Normal coronaries



## **Imaging Studies**

	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
ST segment elevation	YES	no	no
Decreased LV function	no	no	YES/no
Coronary dilation	no	no	YES
Cardiac MRI gadolinium enhancement	YES	?	?
Cardiac catheterization	Normal coronaries	?	Possible coronary artery dilation



## **Imaging Studies**

	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
ST segment elevation	YES	no	no
Decreased LV function	no	no	YES/no
Coronary dilation	no	no	YES
Cardiac MRI gadolinium enhancement	YES	?	?
Cardiac catheterization	Normal coronaries	?	Possible coronary artery dilation



# Comparison of Myocarditis to COVID-19 Related Illnesses

	Vaccine related myocarditis	Acute COVID infection	Multisystem Inflammatory Syndrome in Children
Chest pain	YES	no	no
Symptoms within a few days from mRNA vaccine	YES	no	no
CK-MB	YES	no	maybe
Troponin	YES	no	YES (20-60%)
SARS-CoV-2 IgG	no	maybe	YES
ST segment elevation	YES	no	no
Cardiac MRI gadolinium enhancement	YES	?	?



#### Differential Diagnosis

- Chest Pain
  - Too vast to consider
  - Q: recent (within 5 days) Pfizer BioNTech or Moderna vaccine
  - A: obtain ECG and labs: troponin and CK-MB
- ST segment elevation, troponin and CK-MB elevation with chest pain
  - Myocardial infarction
  - Acute myocarditis
  - Coronary spasm (angina)
  - Q: Kawasaki disease, familial hypercholesterolemia, 10+ year h/o smoking
  - A: myocarditis



#### Differential Diagnosis of Myocarditis

#### Acute myocarditis

- Fulminant, focal, viral, MIS-C
- Q: viral prodrome, h/o COVID infection, cardiac function by echocardiogram
- A: if echo normal and no h/o recent viral illness, it is likely focal myocarditis due to COVID-19 vaccine

#### Consideration, supportive tests

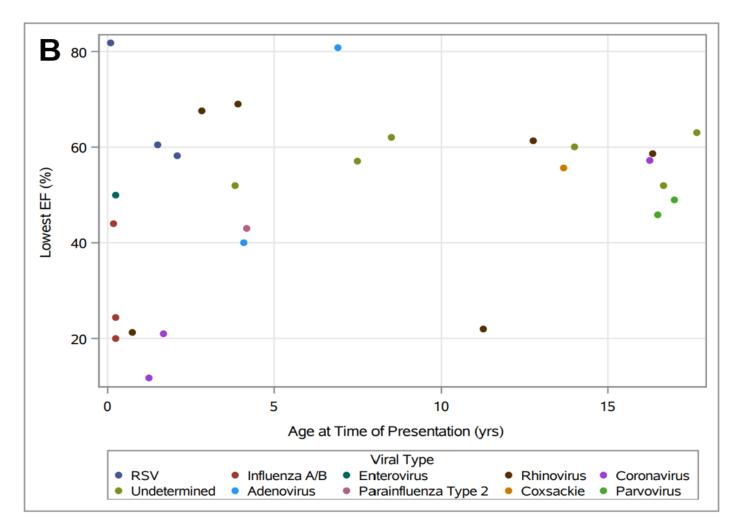
- Possible viral myocarditis: PCR for typical viruses
- Possible reactive myocarditis (MIS-C): SARS-CoV-2 nucleocapsid IgG
- Fulminant myocarditis: monitor vital signs and function
- Trend cardiac enzymes: CK-MB!



#### Viral Myocarditis

Viral etiology across pediatric agerange

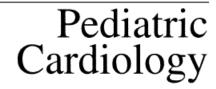
Common in adolescents: Rhino Parvo Coxsackie Corona





#### Acute Myocarditis in Adolescents

Pediatr Cardiol 26:627–631, 2005 DOI: 10.1007/s00246-004-0864-5



© Springer Science+Business Media, Inc. 2005

"Myocardial Infarction" in Adolescents: Do We Have the Correct Diagnosis?

A. Desai, S. Patel, W. Book 1,2

<sup>1</sup>Division of Cardiology, Department of Internal Medicine, Emory University School of Medicine, Atlanta, GA 30322, USA <sup>2</sup>Emory Adult Congenital Cardiac Program, Emory University Hospital, 1364 Clifton Road NE Suite F508, Atlanta, GA 30322, USA



#### Acute Myocarditis in Adolescents

Table 1. Chest pain characteristics, cardiac enzymes, and ECG abnormalities

Mean age (years)	16.7
% male	89
Viral prodrome (%)	44
Mean duration of chest pain (hours)	40
% with ST elevation on initial ECG	89
Peak CK-MB, mean (ng/ml)	65.7
Peak troponin I, mean (ng/ml)	14.0
Coronary occlusion at catheterization (%)	0
Coronary distribution of ECG changes	
Anterior (%)	33
Inferior or inferiolateral (%)	67



#### Focal Myocarditis

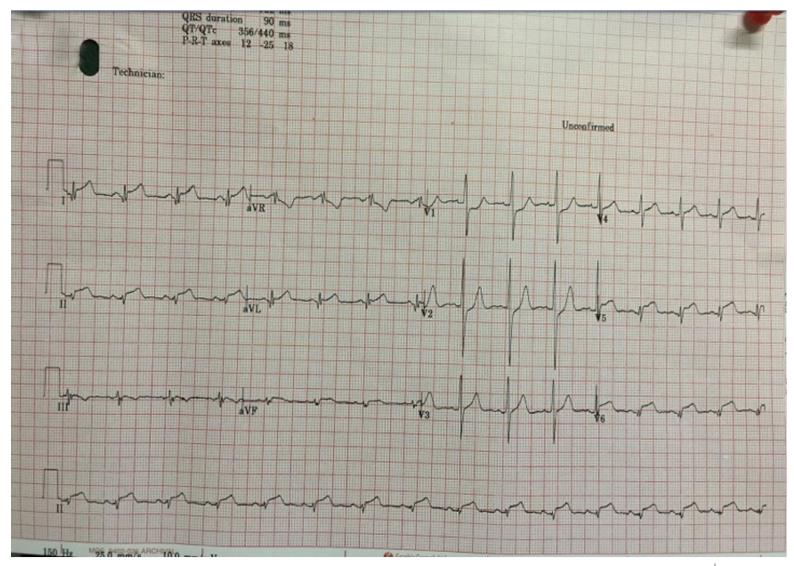
#### **Acute Myocarditis and ST-Segment Elevation**

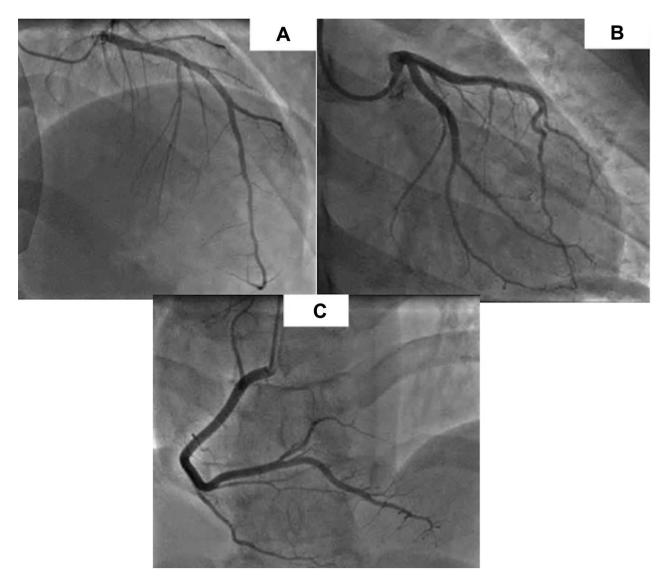


Muhammad Shahid, MRCP, Edward Hoey, MRCP, FRCR, and Sandeep Basavarajaiah, MD, MRCP\*

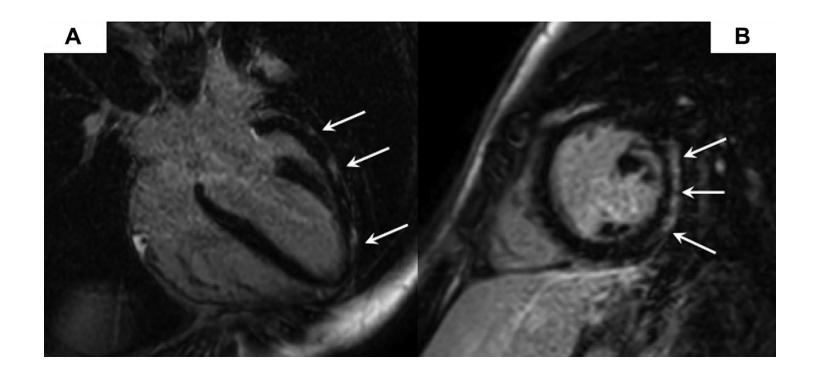
We report a case of focal myocarditis in a young boy mimicking acute ST-segment elevation MI. He presented with chest pain and the EKG changes were consistent with infero-laeral ST-segment elevation MI. Coronary angiogram revealed smooth arteries with no obstruction. Troponin was significantly elevated and the echocardiogram exhibited mildly impaired LV function with hypokinetic inferior and lateral walls. Subsequently performed cardiac magnetic resonance imaging confirmed the diagnosis of myocarditis by exhibiting classic features of delayed gadolinium enhancement in the epi and mid-myocardial regions of the lateral wall sparing the sub-endocardial region. This case exhibits the use of cardiac magnetic resonance imaging for diagnosis in such scenarios as often if the angiogram is normal other differential diagnosis are often speculated without actual evidence © 2016 Elsevier Inc. All rights reserved. (Am J Cardiol 2016;118:1605–1608)







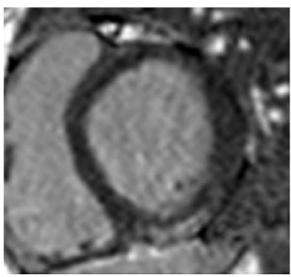
### Focal myocarditis

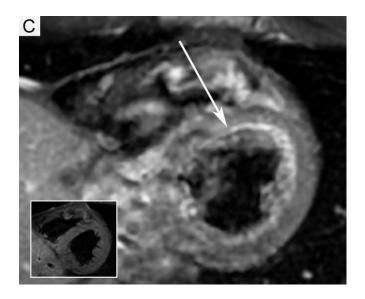




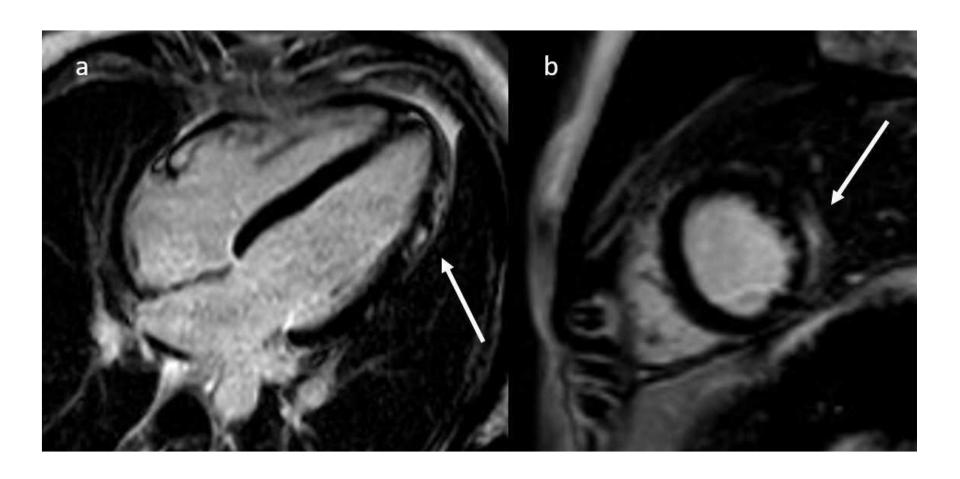
### Focal Myocarditis Imaging











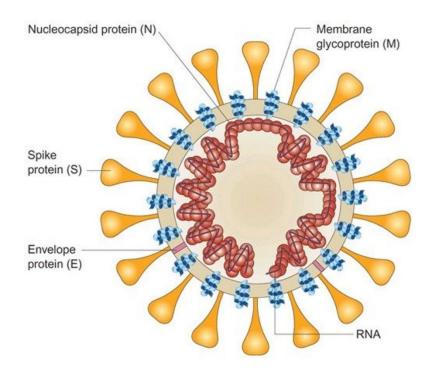


#### Hypothetical Pathomechanism



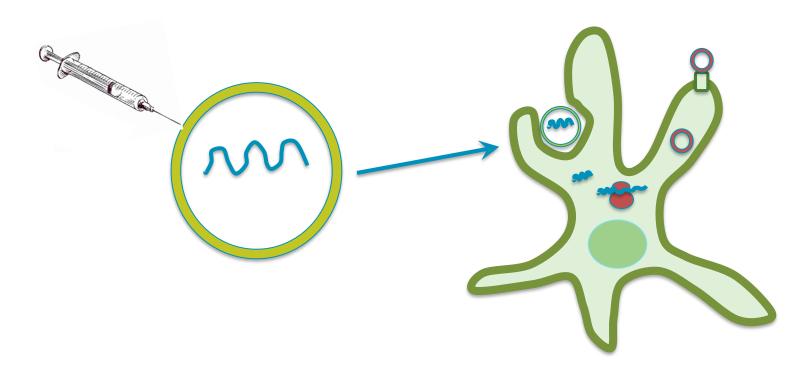


#### Anatomy of the SARS-CoV-2

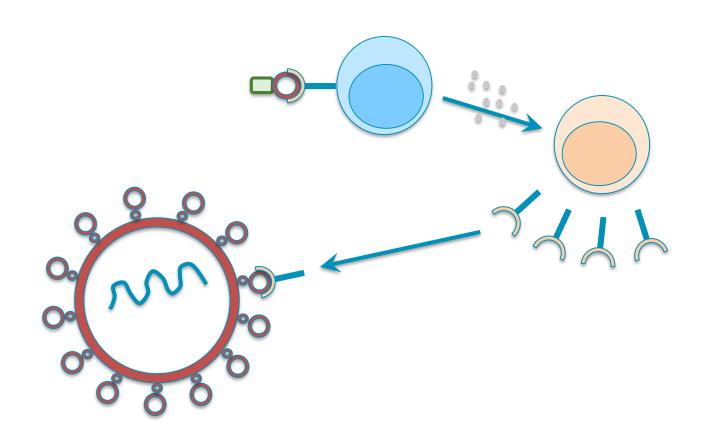




#### Hypothetical Pathomechanism

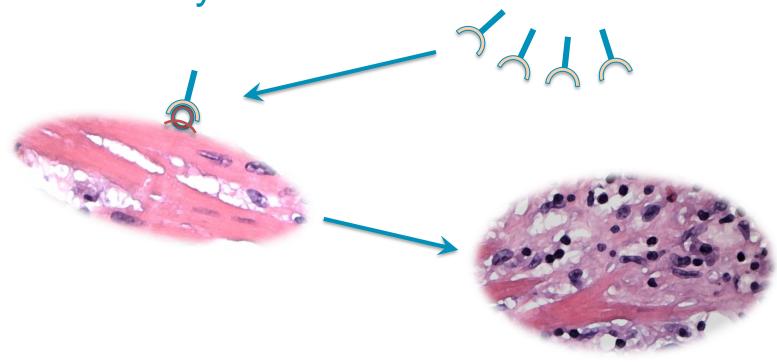








Hypothetical pathomechanism of vaccine associated myocarditis



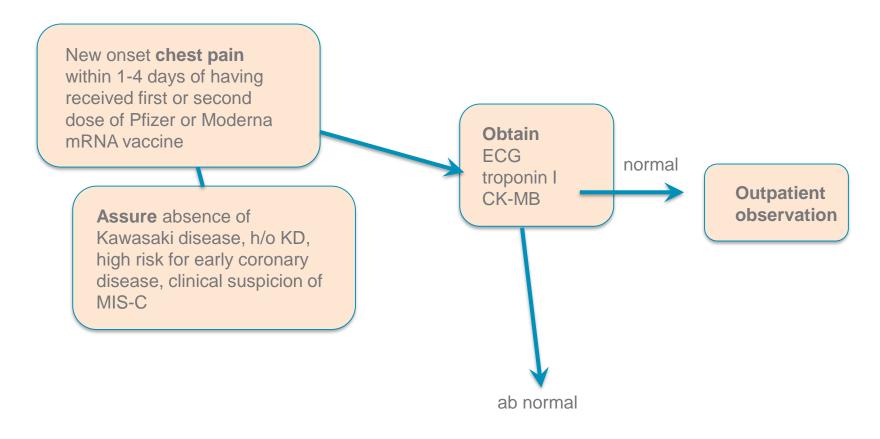


#### What to expect with focal myocarditis

- Chest pain
- Abnormal ECG
- Troponin and CK-MB elevation
- Normal echocardiogram
- Resolution in 2-5 days
- No ECMO
- No long-term sequalae (?)

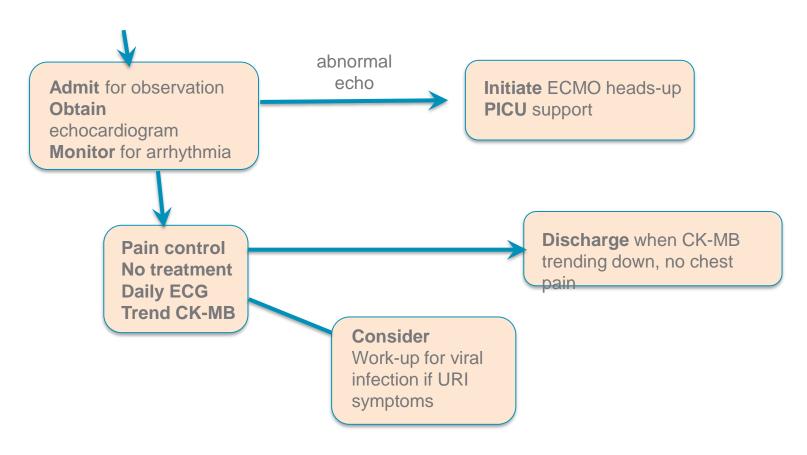


# Proposed algorithm to assess vaccine related myocarditis





# Proposed algorithm to treat vaccine related myocarditis

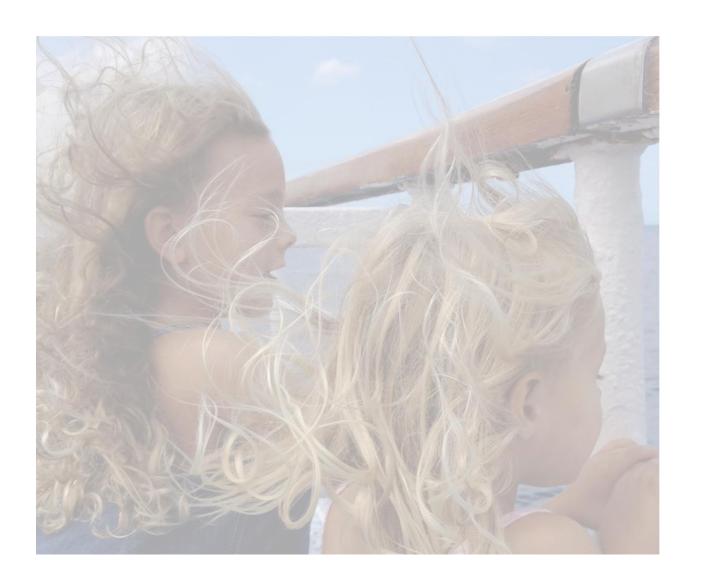




#### Summary

- COVID-19 mRNA vaccine can, rarely, cause myocarditis (1:5000)
- The myocarditis presents with chest pain, ST segment elevation and troponin elevation
- There is no indication for detailed viral PCR testing, unless the symptoms and presentation warrant it
- The myocarditis is focal, self limited, does not require medication
- Currently there is no known long-term sequelae





Thank you for your attention!



#### References

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  Authorizes Pfizer-BioNTech COVID-19 Vaccine for Emergency Use in Adolescents in Another Important
  Action in Fight Against Pandemic. U.S. Food and Drug Administration, FDA, 10 May 2021.
- Wallace M, Woodworth KR, Gargano JW, et al. The Advisory Committee on Immunization Practices' Interim Recommendation for Use of Pfizer-BioNTech COVID-19 Vaccine in Adolescents Aged 12–15 Years — United States, May 2021. MMWR Morb Mortal Wkly Rep 2021;70:749–752.
- "Myocardial infarction in adolescents: Do we have the right diagnosis? Desai et al., Pediatric Cardiology, 2005.
- Acute myocarditis and ST segment elevation. Shahid et al., American Journal of Cardiology, 2016.
- Symptomatic acute myocarditis in seven adolescents following the Pfizer-BioNTech COVID-19 vaccine.
   Marshall et al., Pediatrics, 2021.
- Self-limited myocarditis presenting with chest pain and ST segment elevation in adolescents after vaccination with the BNT162b2 mRNA vaccine. Park et al, Cardiology in the Young, 2021.



#### Q&A



#### **Next Webinar:**

# HHP Care Model and Disease Management Webinar:

GI – Dr. Timothy Swindoll

Thursday, June 24, 2021 5:30pm – 6:30 pm

Please note: webinar topic is subject to change



## Thank you!

- A recording of the meeting will be available afterwards.
- Unanswered question?
  - Contact us at <u>Covid19Bulletin@hawaiipacifichealth.org</u>

