Behavioral Health ECHO

November 30, 2021

Long Term Manifestations of COVID Disease

Dominic Chow, MD, PhD, MPH

Professor John A. Burns School of Medicine





No financial disclosures



Objectives

- Magnitude of Post Acute Sequelae of COVID-19 (PASC) or "Long COVID"
- Symptoms
- Definition
- Possible mechanisms
- Appreciate the pulmonary, cardiac, neurologic, and psychiatric complications
- Potential treatments
- Public health implications



COVID-19 Timeline

- December 31, 2019 ~ clusters of pneumonia reported in Wuhan, China
 - January 11, 2020 ~ first death in China reported
 - January 21, 2020 ~ first US case reported
 - ~ Hawaii 's first case of COVID-19, cruise to Mexico on Grand Princess ship
 - ~ Hawaii's first death, Oahu man who traveled to Las Vegas
 - ~ First mention of Omicron
 - November 28, 2021 ~ Hawaii achieves 72.5% full covid vaccination and 85.4% at least one dose

	Global	U.S.	Hawai'l (HiEMA data)
Confirmed Cases Nov 28, 2021	259,502,031	47,802,459	87,514
Deaths Nov 28, 2021	5,183,003	771,529	1,018
New Cases Nov 28, 2021	611,528	100,455	169



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March 06, 2020

March 31, 2020

November 24, 2021

https://covid19.who.int/ https://health.hawaii.gov/coronavirusdisease2019/

Possible 5th COVID Surge in the United States



Daily Trends in Number of Cases and 7-Day Cumulative Incidence Rate of COVID-19 Cases in The United States Reported to



10/16/20 - 105 (105/911 = 11.5%) patients results as positive/detected.

Count of Ethnicity		10/16/2020
Ethnicity	Total	
Arab/Arabian (Middle East)	1	1%
Black/Other	1	1%
Chinese	1	1%
Chuukese	47	45%
Declined	1	1%
Filipino	4	4%
Japanese	2	2%
Japanese/Okinawan	1	1%
Korean	3	3%
Laotian	1	1%
Marshallese	6	6%
Mexican	1	1%
Other Micronesian	3	3%
Part Hawaiian	6	6%
Pohnpeian	2	2%
Puerto Rican	1	1%
Samoan	14	13%
Thai	1	1%
Tongan	1	1%
Vietnamese	1	1%
White/Caucasian	7	7%
Grand Total	105	

Queen Emma Clinics

Count of ETHNICITY		12/31/2020
ETHNICITY	Total	
Arab/Arabian (Middle East)	3	2.2%
Black/Other Chinese	1	0.7% 0.7%
Chuukese	47	34.8%
Declinea	1	0.7%
Filipino	6	4.4%
Japanese	4	3.0%
Japanese/Okinawan	1	0.7%
Korean	3	2.2%
Laotian	1	0.7%
Marshallese	12	8.9%
Mexican	1	0.7%
Other Micronesian	5	3.7%
Part Hawaiian	10	7.4%
Pohnpeian	3	2.2%
Puerto Rican	1	0.7%
Samoan	21	15.6%
Thai	1	0.7%
Tongan	1	0.7%
Vietnamese	3	2.2%
White/Caucasian	9	6.7%
Grand Total	135	

Count of ETHNICITY			3/2/2021	
ETHNICITY	-	Total		
Arab/Arabian (Middle East	st)	3	2.1%	
Black/Other		1	0.7%	
Chinese		1	0.7%	
Chuukese		47	33.3%	
Declined		1	0.7%	
Filipino		7	5.0%	
Japanese		4	2.8%	
Japanese/Okinawan		1	0.7%	
Korean		3	2.1%	
Laotian		1	0.7%	
Marshallese		12	8.5%	
Mexican		1	0.7%	
Other Micronesian		5	3.5%	
Part Hawaiian		8	5.7%	
Pohnpeian		3	2.1%	
Puerto Rican		1	0.7%	
Samoan		24	17.0%	
Thai		1	0.7%	
Tongan		1	0.7%	
Vietnamese		5	3.5%	
White/Caucasian		10	7.1%	
Portuguese		1	0.7%	
Grand Total		141	1416	10%





Footnotes: Based on first non-White race listed; **White with no other race listed; †Native Hawaiian as any listed race. * Excludes residents diagnosed out-of-state (n=1,010), non-residents (n=904), and cases with no specific race information available (n=7,134); Hospitalization status is unknown for 2,544 cases with known race information. Horizontal bands indicate 95% confidence interval for Case, Hospitalization, or Death %

Pacific Islanders and Filipinos are disproportionately effected.

Hawai'i's Micronesian migrants get limited access to health care

The Next Epidemic...

Post Acute Sequelae of Sars-COV-2 (PASC) – "Long COVID"

- 10-40% of all COVID infections
- Duration unknown, possibly permanent
- High degree of disability

7 Celebrities With Scary Long-Term COVID Symptoms THESE STARS ARE STILL STRUGGLING TO MANAGE TERRIFYING SYMPTOMS, MONTHS AFTER TES











"I had swollen hands and feet, an unceasing migraine and fatigue that limited my every move...



Hawaii state Sen. Kalani English stepping down due to long-term COVID-19 symptoms

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By Sophie Cocke · April 27, 2021

Senale Matority Leader J. Kalani English spoke loday in the Capitol rolunda.





Lung Disease & Respiratory Health > Coronavirus > News

WEBMD NEWS BRIEF

More Than 100 Million People Worldwide Have or Had Long COVID: Study By Carolyn Crist



Nov. 18, 2021 -- More than 40% of COVID-19 survivors across the world have or had long-term effects after recovering according to a new study by recearchers at

Coronavirus Outbreak



Symptoms of Long COVID

Top symptoms

- Fatigue
- Dyspnea
- Anxiety
- Concentration / cognitive changes

People hospitalised during acute phase of COVID-19 Based on 16 studies with 3172 people aged 18 or over



Post-traumatic stress disorder Sleep disorder Impaired mobility Care dependency Reduced quality of life psychological symptoms

Upper respiratory Voice change Nasal congestion*

Cardiopulmonary Breathlessness Palpitations*

Urinary continence

Acute COVID-19 Post-acute COVID-19

Subacute/ongoing COVID-19

Chronic/post-COVID-19



Before symptom onset

Nalbandian A, Nature Medicine volume 27, pages601–615(2021)

W NEWS - CORONAVIRUS NEWS

WHO Releases First Official Long COVID Definition

By Carla Delgado | Published on October 25, 2021 NEW



S Fact checked by <u>Nick Blackmer</u>



FG Trade / Getty Images

Key Takeaways

- Long COVID has been used to refer to a range of new, returning, or ongoing symptoms after initial COVID-19 infection.
- This month, the WHO has published an official definition of post-COVID-19 to advance research and help diagnose individuals.
- The definition can change and be refined further as new evidence emerges regarding the condition.

A clinical case definition of post COVID-19 condition by a Delphi consensus

6 October 2021

Adventisement





Table 3. A definition of post COVID-19 condition

Post COVID-19 condition occurs in individuals with a history of probable or confirmed SARS-CoV-2 infection, usually 3 months from the onset of COVID-19 with symptoms that last for at least 2 months and cannot be explained by an alternative diagnosis. Common symptoms include fatigue, shortness of breath, cognitive dysfunction but also others* and generally have an impact on everyday functioning. Symptoms may be new onset following initial recovery from an acute COVID-19 episode or persist from the initial illness. Symptoms may also fluctuate or relapse over time.

A separate definition may be applicable for children.





Pathophysiology of shared multiorgan injury



Gupta A et al. Nat Med . 2020 Jul;26(7):1017-1032.

Potential Mechanisms of Long COVID

- Autoantibodies
 - Woodruff M, medRxiv, 2021.09.21.21263845
- Post viral transcripts
- Viral latency
- Monocyte activation
 - Patterson BK, bioRxiv, 2021.
- Inflammation
 - Peluso MJ, Cell Rep. 2021 Aug 10;36(6):109518.
- Micro clots
 - Pretorius E, Cardiovasc Diabetol 20, 172 (2021).



Cardiovascular Complications

- Myocarditis
- Cardiomyopathy
- Myocardial infarction
- Heart failure
- Arrhythmias
- Cardiac arrest
- Thromboembolic events

Poor prognosis

- Increased troponin
- Increased BNP (brain natriuretic peptides)

RESEARCH LETTER

Cardiovascular Magnetic Resonance Findings in Competitive Athletes Recovering From COVID-19 Infection



- 26 competitive college athletes diagnosed with COVID-19 (RT-PCR)
- None were hospitalized
- Majority did not report symptoms
- 12 (46%) had evidence of myocarditis or prior myocardial injury by cardiac magnetic resonance imaging routinely performed for positive testing results (range, 12-53 days)



A CT scan of the chest of a 66-year-old male reveals patchy rounded hazy spots throughout the lungs. He had tested positiv for the coronavirus and experienced shortness of breath. Steven Needell/Science Source

Radiology: Cardiothoracic Imaging

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Original Research	🔂 Free Access

Chest CT Findings in Cases from the Cruise Ship "Diamond Princess" with Coronavirus Disease 2019 (COVID-19)

🐵 Shohei Inui 🖂 🕪 Akira Fujikawa, 🐵 Motoyuki Jitsu, Naoaki Kunishima, 🕪 Sadahiro Watanabe, Yuhi Suzuki, 🛈 Satoshi Umeda, Yasuhide Uwabe

✓ Author Affiliations

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Pulmonary Sequela Common

- Asymptomatic cases showed more ground glass opacity (GGO) over consolidation (83%), while symptomatic cases more frequently showed consolidation over GGO (41%).
- This study documented a high incidence of subclinical CT changes in cases with COVID-19. Compared to symptomatic cases, asymptomatic cases showed more GGO over consolidation and milder extension of disease on CT.

Neurological Complications

- Headache
- Seizure
- Ischemic and hemorrhagic strokes, including large vessel stroke in young people
- Corticospinal tract involvement
- Cognitive dysfunction
- Olfactory and gustatory dysfunctions (can present as first or sole symptom)
- Acute encephalitis, necrotizing encephalopathy, disseminated encephalomyelitis, transverse myelitis
- Guillain-Barre syndrome

risks for long term cognitive impairment and neurodegenerative diseases

Psychiatric Complications

- Depression
- Anxiety
- Fatigue
- Post-traumatic stress disorder
- Dysexecutive syndrome (brain fog)

Long Haulers

I used to go to the gym three times a week. [Now,] my physical activity is bed to couch, maybe couch to kitchen.

Athena Akrami, University College London

Rogers et al. Lancet Psychiatry. 2020 Jul; 7(7): 611-627

Thromboembolic Complications

- Pulmonary embolism
- Peripheral deep vein thrombosis
- Disseminated intravascular coagulation
- Ischemic strokes

cumulative incidence is 20-30% across several studies

Middeldorp. J Thromb Haemost. 2020 Aug;18:1995-2002 Flok. Thromb Res. 2020 Jul;191:145-147 Lodigiani. Thromb Res. 2020 Jul;191:9-14

Long COVID in Primary Care

- Highly variable
- No current guidelines on treatment
- Long term disability



'Long-COVID': a cross-sectional study of persisting symptoms, biomarker and imaging abnormalities following hospitalisation for COVID-19

 In those discharged with elevated biomarkers, 30.1% and 9.5% had persistently elevated d-dimer and C reactive protein, respectively. 38% of chest radiographs remained abnormal with 9% deteriorating.

Thorax. 2021 Apr;76(4):396-398. doi: 10.1136/thoraxjnl-2020-215818.

Post COVID Care

Pulmonary/cardiovascular



Nalbandian A, Nature Medicine volume 27, pages601–615(2021)

the**bmj** Visual summary 🐠

Post-acute covid-19 appears to be a multi-system disease, sometimes occurring after a relatively mild acute illness. Clinical management requires a whole-patient perspective. This graphic summarises the assessment and initial management of patients with delayed recovery from an episode of covid-19 that was managed in the community or in a standard hospital ward.

An uncertain picture



The long term course of covid-19 is unknown. This graphic presesents an approach based on evidence available at the time of publication.

However, caution is advised, as patients may present atypically, and new treatments are likely to emerge

Managing comorbidities

Many patients have comorbidities including diabetes, hypertension, kidney disease or ischaemic heart disease. These need to be managed in conjuntion with covid-19 treatment. Refer to condition specific guidance, available in the associated article by Greenhalgh and colleagues

"Long covid" in primary care

Assessment and initial management of patients with continuing symptoms



Investigations

Clinical testing is not always needed, but can help to pinpoint causes of contiuing symptoms, and to exclude conditions like pulmonary embolism or myocarditis. Examples are provided below:

Blood tests

Full blood count	Electrolytes
Liver and renal fund	tion Troponin
C reactive protein	Creatine kinase
D-dimer Brain na	atriuretic peptides
Ferritin – to asses and pro	s inflammatory thrombotic states

Other investigations

Chest x ray Urine tests

12 lead electrocardiogram

Prolonged covid-19 may limit the ability to engage in work and family activities. Patients may have experienced family bereavements as well as job losses and consequent financial stress and food poverty. See the associated while her Considerable and



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The Queen's Medical Center Post COVID Care Paradigm



Local Studies at Queens

- COVID Seroprevalence Study
 - Myra W. and Jean Kent Angus Foundation
- Telmisartan Inpatient / Ambulatory
- RECOVER COHORT NETWORK

Actual Patient Visit

- •<u>Clinical Portion</u>
- Patient registers
- Vital conducted (temperature, BP, Pulse ox, Resp Rate, HT and wt), PQS4 for anxiety and depression
- COVID Questionnaire
- International Physical Activity Questionnaire
- Beck's Depression Questionnaire
- Hamilton Anxiety Questionnaire
- MOCA
- Frailty Questionnaire
- Physical examination
- NIH Stroke Scale
- Regular neuro test vibration, DTR's, strength
- 6 minute walk test

Actual Patient Visit

Procedures / Tests

- Order the following procedures:
 - CXR
 - PFT
 - ECG
 - ECHO with cardiac strain
- Patient obtain blood test
 - CBC with diff to look at % monocytes and lymphocytes
 - Chem to look at renal and liver function
 - COVID antibody quantification
 - 5th generation troponin
 - Inflammatory markers IL6, D-Dimer, fibrin/fibrinogen, procalcitonin
 - Thrombotic markers

Long COVID Treatment

• Early in the disease course – No clinical guidelines • May provide insight into complications in the future • Cardiovascular - Referral to Cardiology • Primary / secondary prevention • Pulmonary – Referral to Pulmonary group • Use of anticoagulants in thrombi • Neuro/psych – Referral to Neurology and Psychiatry • Cog Rehab • Other – GI, Rheumatology and Nephrology Services • May help in getting your patients into seeing the subspecialists more quickly • May help in coordinating care with PT/OT, social services • Clinical trials

Current Therapies

Fatigue / Chronic fatigue syndrome

- Exercise target of 80% Max heart rate
- Improve sleep
- Get vaccinated with COVID vaccine
- SSNI
- Counseling
- Aspirin and Statin

Unclear use

- Hydroxychloroquine
- Ivermectin
- Maraviroc
- Iodine, high doses of vitamin A,D, E and hydrogen peroxide

Current Therapies - Pulmonary disease

- Exercise / deep breathing
- Oxygen supplementation of <88% RA or drop of 3% from baseline
- ICS/LABA
- ACEI/ARB
- Oral steroids Following SARS-CoV-2 pneumonitis, a cohort of patients are left with both radiological inflammatory lung disease and persistent physiological and functional deficit. Early treatment with corticosteroids was well tolerated and associated with rapid and significant improvement.*
- Herbal Supplements
 - Allium sativum, Althaea officinalis, Andrographis paniculate, Commiphora molmol, Cymbopogon citratus, Echinacea sp, Eucalyptus globulus, Hedera helix, Justicia pectoralis, Magnolia officinalis, Malva sylvestris, Mikania glomerata, Pelargonium sidoides, Pimpinella anisum, Platycodon chinensis, Polygala senega, Scutellaria baicalensis, Silybum marianum, Thymus vulgaris, Cinchona

Ann Am Thorac Soc. 2021 May;18(5):799-806.

Telmisartan Studies

 Fewer Deaths in Hospitalized COVID Patients on an ACEI or ARB. MedRxiv.org.

Current therapies

Brain Fog

- Cog Rehab, reminders, supportive therapy
- Stimulants

Anxiety

- Counseling
- Zoloft or SSRI's
- Benzodiazepine
- Medical Marijuana / CBD

Current therapies

Postural orthostatic tachycardia syndrome (POTS)

- Exercise on recumbent bicycle
- Oral rehydration
- Increase salt intake
- Mineralocorticoids/ BB/ Midodrine

Thrombotic events / Endotheliitis

- NOAC
- Aspirin
- ? Aphersis (plasma exchange)

Current therapies

Loss of smell and taste

- Cog Rehab using essential oils and foods
- Flonase
- Dymista

Herbal remedies that are not recommended

- Ginkgo (Ginkgo biloba L., Ginkgoaceae), St. John's wort (Hypericum perforatum L., Hypericaceae), and valerian (Valeriana officinalis L., Caprifoliaceae) because they may increase the effects of sleep agents. They may also cause irregular heart rhythms.
- Ginseng (Ginseng sp., Araliaceae), licorice (Glycyrrhiza glabra L.), and milk thistle [Silybum marianum (L.) Gaertn.], because they may cause high blood pressure and a rapid heart rate.
- Garlic (Allium sativum L., Amaryllidaceae), ginkgo, green tea [Camellia sinensis (L.) Kuntz., Theaceae], feverfew (Tanacetum parthenium L., Asteraceae), ginger (Zingiber officinale L.) and Saw palmetto [Serenoa repens (W. Bartram) Small, Arecaceae], because they may cause prolonged bleeding.
- • Garlic, in addition, can increase the effects of some OTC pain relievers.
- Ephedra: Several studies and clinical trials have been carried out to identify drugs that can effectively treat the disease, but, at the moment, the strategies to deal with the infection are only supportive

Potential Therapies in the future

• Antivirals

- Pfzier PAXLOVID
- Merck Molnupiravir
- Mast cell inhibitors: Covid-19 hyperinflammation and post-Covid-19 illness may be rooted in mast cell activation syndrome*
- Inhaled Telmisartan

.Afrin LB, Weinstock LB, Molderings GJ. Int J Infect Dis. 2020 Nov;100:327-332.



COVID-19 Community Ambulatory Recovery and Evaluation (CARE) Center





Research Dr. Todd Seto / QMC Director, Academic Affairs and Research





THE QUEEN'S HEALTH SYSTEMS



The Queen's Health Systems COVID-19 Community Ambulatory Recovery and Evaluation (CARE) Program provides specialized after-care to patients diagnosed with COVID-19 and are in recovery.

As part of the program, the Post-COVID CARE Clinic provides QHS and community patients with diagnostic services and complete cardiovascular, pulmonary, and neuropsychiatric evaluation.

COVID-19 may increase the risk of longterm health problems, prolonged illness, and persistent symptoms—even in young adults and persons with no underlying medical conditions or those who were not hospitalized.

Queen Emma Clinics

- Queen Emma Tower, Ground Floor 1301 Punchbowl Street, Honolulu, HI 96813
- 😢 (808) 691-4970
 -] www.covid.queens.org/clinic

Patients recovering from COVID-19 often experience symptoms that can indicate long-lasting side effects to body systems and organs, including:

Heart

- Damage to heart muscle
- Heart failure

Lungs

- Pulmonary disease/embolism
- Damage to lung tissue
- Restrictive lung failure

Brain and Nervous System

- Stroke
- Loss of memory/ability to concentrate
- Loss of sense of smell

Mental Health

- Anxiety and/or depression
- Post-traumatic stress disorder
- Sleep disturbance

Joints and Muscles

- Pain or weakness in joints and muscles
- Chronic fatigue
- Loss of stamina



If you are a patient experiencing post-COVID symptoms, or a provider who would like to refer a patient, please contact us at (808) 691-4970 or visit www.covid.queens.org/clinic to learn more about the clinic and download the Patient Referral Form.



Post COVID Recovery and Care Clinic

Dedicated to the treatment and management of patients who are suffering from complications stemming from their COVID infection.

Post COVID Recovery Referral and Questions Call: (808) 691-4970

Download Referral Form

The Queen Emma Clinic Post-COVID Eligibility Criteria:

- Patient must have a confirmed positive COVID-19 infection diagnosis.
- Patient must be at least 10 days' post COVID infection with no current signs and symptoms of active infection (i.e. – fever ≥ 100.4° F or chills, cough, shortness of breath or difficulty breathing, etc.)
- Patient referral from assigned PCP.
- PCP may refer patient via Post COVID care referral form.

https://covid.queens.org/post_covid_clinic/



THE QUEEN'S HEALTH SYSTEMS COVID-19 COMMUNITY AMBULATORY RECOVERY AND EVALUATION

The Queen Emma Clinic Post-COVID Bigibility Criteria:

- Patient must have a confirmed positive COVID-19 infection diagnosis.
- Patient must be at least 10 days' post COVID infection with no current signs and symptoms of active infection (i.e. - fever ≥ 100.4° F or chills, cough, shortness of breath or difficulty breathing, etc.)

o NoFault

o Seoul Medical

- Patient referral from assigned PCP.
- PCP may refer patient/via Post COVID care referral form.
 - QEC accepts most insurances except the following:
 - o Medicare HMO
 - o Commercial HMO

o Kaiserhealth plans

Group (SMG)

o Tricare Prime o Worker's Compensation

Referring Provider:

CD-10 code (s):		
		
a tient contact information:		
ame:	DOB:	
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If you have any questions, please feel free to call us at (808) 691-4970.

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MAHALO!

