FADONA 2023

Restorix Health

"Latest Updates in COVID Skin and Other Long-Hauler Issues

Presenter

Pamela Scarborough PT, DPT, CWS, FAAWC Vice President of Clinical Affairs Restorix Health

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OBJECTIVES

- At the end of this presentation, participants should be able to:
- Describe multiple and varied skin manifestations related to COVID-19;
 Relate some of the COVID-19 skin disruptions to other skin manifestations that look similar to, but are NOT COVID-19 skin issues;
- 3. Describe COVID-19 disease affect on wound healing
- Define Post-Acute Sequelae of SARS CoV-2 infection and the proposed mechanisms leading to Long COVID
- Describe the theory of why Post-Acute Sequalae of SARS-CoV-2 (PASC) develops in some people
- List proposed treatment and rehabilitation considerations for patients/residents with

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COVID-19 EPIDEMIC

- COVID-19 pandemic caused by SARS-CoV-2
 - Primarily triggers respiratory tract infections
 - > Affects upper or lower respiratory tracts
 - Spreads same way other coronaviruses do
 - Mainly through person-to-person contact
 - Infections range from mild, moderate to severe to deadly outcomes
- Originally thought to be only respiratory disease
- Current research demonstrates significant extrapulmonary involvement
- New variants appear to cause less pulmonary involvement for most people

CYTOKINE STORM & COVID-19



- Cytokines are part of immune system
 Causes acute hyperinflammatory response
- Immune cells spread beyond infected body parts
- Inflammatory response to infection
- Attacks healthy tissues
- Causes blood clots
 - Coagulopathy
 - Creates decreased blood flow to organs
- Skin is largest organ
- Blood flow and inflammatory processes often manifest on skin and mucous membranes

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CURRENT MOST COMMON SYMPTOMS OF XBB.1.5 OMICRON VARIANT

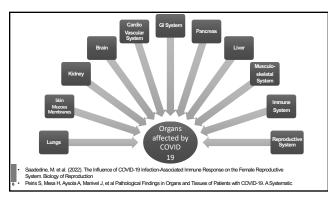
Fatigue

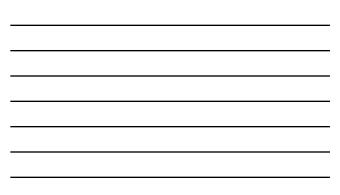
- Congestion
- Body or muscle aches
 - Mild symptoms
 - Asymptomatic



- Cough
- Headache
- Nausea/vomiting

Silent symptoms of COVID-19 include skin and mucocutaneous symptoms





CURRENT PRIMARY STRAIN OF COVID-19 Subvariant XBB and XBB.1.5



- Labeled globally as Omnicron family subvariant currently responsible for bulk of new COVID-19 cases since October 2022
- Combination of 2 different variants BA.2.10.1 and BA.2.75
- Note: More than 50 COVID-19 variants across the globe
 Most contagious Omicron mutation to date WHO & CDC
- XBB.1.5 termed "Kraken" strain thought to be 5 times more contagious than earlier Omicron strains
- > Has characteristics that allow it to spread easier among both vaccinated and recently COVID-19-recovered individuals
- Common symptoms associated with XBB.1.5 strain can be easily
- dismissed as seasonal illness or flu
- https://www.yalemedicine.org/news/omicron-xbb-kraken-subvariant

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V	VHO label	Lineage #	US Class	%Total	95%PI	
	Omicron	XBB.1.5	VOC 90	.2%)	87.4-92.4%	
		BQ.1.1	VOC 3.5	5%	2.5-4.9%	
_		XBB	VOC 2.5	5%	1.5-4.1%	
_		onsible fo Infectio /covid.cdc.	CDC B.1.5. Va or 90% of N ons throug gov/covid-da proportion ccessed 3/19	New C h 3/18 ata-trac s	8/23	

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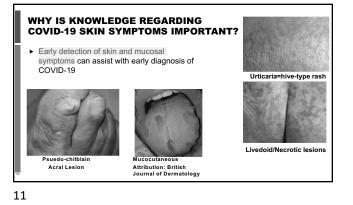
INITIAL COVID SKIN MANIFESTATIONS RECOGNITION IN HEALTHCARE

 April 2020 International League of Dermatological Societies and American Academy of Dermatology established international registry for COVID-19 dermatological manifestations

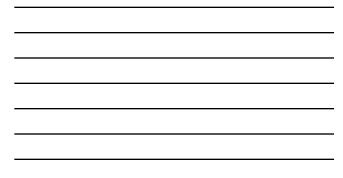


- At the same time, clinicians in LTC facilities reporting skin manifestations that looked like pressure injuries, but were NOT related to pressure
- Also, anecdotally, residents with wounds that HAD been improving began to stall or get worse after surviving COVID-19
- ► April 2020 the journey to learn about COVID skin manifestations began

httos://www.aad.ore/member/oractice/coronavirus/reeistry Mantovani, A., Morrone, M.C., Patrono, C. et al. Long Covid: where we stand and challenges ahead. Cell Death Differ 29, 1891–1900 (2022).



HOW DOES SARS-COV-2 VI	RUS INFECT THE SKIN?
Direct Infection of Endothelial Cells	Indirect Infection of Endothelial Cells
 Small blood vessels in skin are targets for SARS-CoV-2 Lymphocyte skin infiltration in patients with COVID-19 Evidence that virus enters skin through blood vessels after systemic infection 	 Superficial layers of skin prevent viral entry with normal conditions Skin disruptions (e.g., wounds) May allow contamination of underlying tissues
Khezri, Mohammad Rafi; Ghasemnejad-Berenji, Morteza PhD to Multiply and Transmit?, Advances in Skin & Wound Care: (



Endothelial Injury	Endotheliitis
Coagulopathy associated abnormalities	Petechiae Dermal necrosis Dermo-hypodermal/superficial thrombi Deep dermis thrombi
Vasculitis	Livedo Purpura Subcutaneous lymphocytic vasculitis Lymphocytic infiltration of vessels
Possible cytopathic effects	Intranuclear viral inclusions Multinucleated cells Intraepidermal vesicle Dyskeratosis Necrotic keratinocytes

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SKIN RASHES PREDICTIVE SYMPTOM OF COVID-19

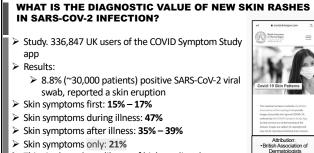
- Virus causes wide variety of skin symptoms
- These skin changes may have diagnostic value for SARS-CoV-2 infections
- Support studies suggest that skin rash may be <u>predictive symptom</u> of COVID-19 infection

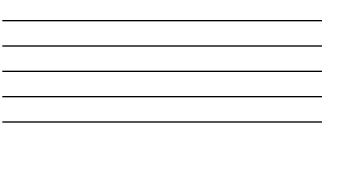


In nine percent of cases, dermatologists identified

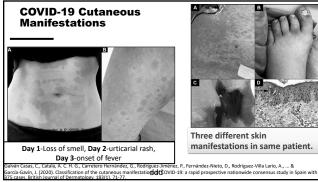
outbreaks of small blisters, commonly itchy, that appeared on the trunk of the body

Bataille V, Visconti A, Rossi N, Murray B et. al. Diagnostic value of skin manifestations of SARS-CoV-2 infection. Young S, Fernandez AP. Skin manifestations of COVID-19. In COVID-19 Curbside Consultants. Cleveland Clinic. Posted May 7, 2020.







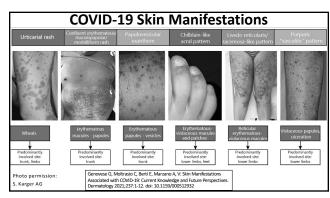


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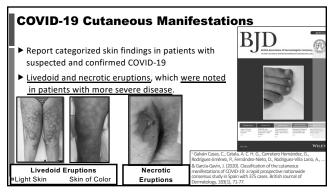
True Incidence of COVID-19 Related Skin Injuries Currently Unknown



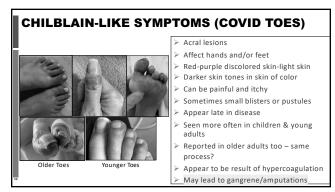
• Many of the skin changes mimic known dermatologic disorders including pressure injuries, Kennedy Terminal Ulcer, and arterial insufficiency wounds

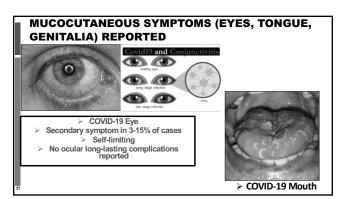


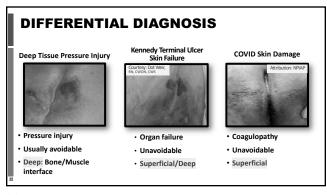








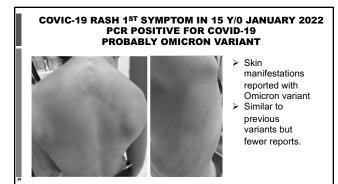




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OMICRON VARIANT SKIN SYMPTOMS

- Overall symptoms described as milder than earlier variants
- Symptoms similar to those of common cold in some people
- Omicron skin manifestations include:
 - Rashes
 - Dry lips
 - Grey/blue-tinged lips or nailbeds



SKIN & MUCOUS MEMBRANE SYMPTOMS REPORTED WITH OMICRON VARIANT

- Chilblain-acral lesions-fingers/toes
- Chapped or sore lips
- Xerostomia (dry mouth)
- Oral lesions
- COVID tongue
- Dry skin
- Other rash-like symptoms



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DOCUMENTATION BY PROVIDERS AND WOUND SPECIALISTS FOR COVID SKIN/WOUND HEALING ISSUES

Dr. Vycki Nalls, PhD, GNP-BC, ACHPN, CWS

- "Wound healing: secondary effects from COVID-19 due to hypoxia, poor nutritional intake, and debility.
- Delayed wound healing expected due to these effects, and it would not be a surprise if the wound does not heal or declines further given patient's declining status."

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ICD-10 DOCUMENTATION FOR DR. NALLS' PATIENT

• L89.150 Pressure ulcer of sacral region, unstageable (HCC 158)

 Unstageable pressure injury to sacrum, with delayed wound healing due to comorbid conditions of hypoxia, poor nutrition, debility, and overall decline from COVID-19 infection.

• U07.1 COVID-19

• COVID positive patient with decline for aggressive management.

D68.8 Other specified coagulation defects (HCC 48)

• Coagulopathy due to COVID-19.

ICD-10 CODES FOR COVID-DERMATOLOGIC MANIFESTATIONS

- > Use U07.1 as first diagnosis for patients with confirmed COVID-19.
- Add an additional diagnosis for pneumonia or other conditions, or symptoms.
- D68.8 is a specific ICD-10 code to indicate a diagnosis of other specified coagulation defect.
 COVID toes/fingers (acral lesions)
- L99 specifies a <u>diagnosis of other disorders of skin and</u> <u>subcutaneous tissue in diseases classified elsewhere (rashes)</u>

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LONG COVID NOMENCLATURE

- ▶ Post-acute sequelae of SARS-CoV-2 (PASC) new formal name
- ► Post-COVID Syndrome (PCS)
- ► Long COVID
- COVID Long Haulers

POST-ACUTE SEQUELAE OF SARS-COV2 INFECTION

- Described by WHO as persistence of symptoms or new symptoms more than 30 days post-SARS-CoV-2 infection
- CDC: 4 or more weeks after infection
- British NIH and Care Excellence (NICE): 12-weeks during or after infection; not explained by alternative diagnosis
- These longer effects of COVID-19 are actively being investigated and defined
- Clinical definition and understanding of underlying mechanisms of Long COVID are still in flux

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POST-ACUTE SEQUELAE OF SARS-COV2 INFECTION

- ► Lingering symptoms may persist months and in some cases years after the acute infection
- ► ICD-10 CM code for "post COVID-19 condition, unspecified"=U09.9
- Deployment of an ICD-10-CM code in US took nearly 2 years <u>after</u> patients began describing their symptoms
- Countries around the world are all dealing with Post COVID in their populations

Pfaff ER, Madlock-Brown C, Baratta JM, et al. Coding Long COVID: Characterizing a new disease through an ICD-10 lens. Preprint. medRxiv. 2022;2022.04.18.22273968. Published 2022 Sep 2. doi:10.1101/2022.04.18.22273968

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POST-ACUTE SEQUELAE OF COVID: FACTS OVERVIEW

- Legacy of acute SARS-CoV-2 infection, affecting 10-69% of patients with different signs and symptoms across a wide range of organs and systems
- Most frequent manifestations of PASC, compromised lung functions, neurocognitive alterations; alterations of cardiovascular functions and increased risk of acute cardiac events; and fatigue.
- SARS-CoV-2 virus seeds and persists in different organs and tissues.
- Pathogenesis of PASC is multifactorial and includes:
 Virus seeding and persistence in different organs; activation and response to unrelated viruses (e.g., EBV); autoimmunity; uncontrolled inflammation.
- Biomarkers of clinical PASC include levels of IgG, cytokines, chemokines, PTX3, and interferons.
- Mantovani, A., Morrone, M.C., Patrono, C. et al. Long Covid: where we stand and challenges ahead. Cell Death Differ 29, 1891–1900 (2022)
 Su; Yuan D, Chen DG, et al. Multiple early factors anticipate post-acute COVID-19 sequelae. Cell. 2022;185(5):881-895.e20. doi:10.1015/cell.2022.01.012

POTENTIAL CONTRIBUTORS TO PASC SYMPTOMS

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approaches may be required to

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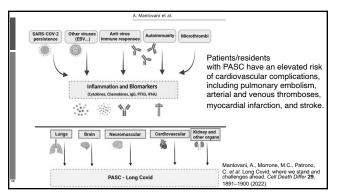
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IP

- Injury to one or multiple organs,
- Persistent reservoirs of SARS-CoV-2 in certain tissues, · Re-activation of neurotrophic pathogens such as The individualized nature of PASC symptoms suggests to herpesviruses under conditions of COVID-19
- immune dysregulation, SARS-CoV-2 interactions with host
- microbiome/virome communities,
- Clotting/coagulation issues,
- Dysfunctional brainstem/vagus nerve signaling,
- Ongoing activity of primed immune cells,
- Autoimmunity due to molecular mimicry between pathogen and host proteins.

Edward Galaid, MD, RSFH Medical Director of Occupational Medicine. Roper St. Francis Healthcare. Management of Post-

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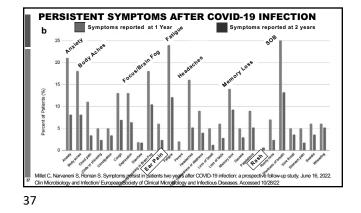


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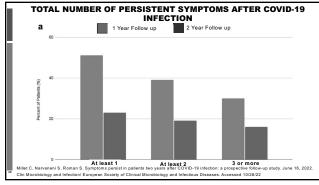
LONG COVID = POST-ACUTE **SEQUELAE OF SARS-COV-2 (PASC)**

- ▶ Research indicated an ongoing, sustained inflammatory response following mild, moderate, and severe SARS-CoV-2 infections
- ▶ "We can show that the macrophages from people with mild COVID-19 exhibit an altered inflammatory and metabolic expression for three to five months post-infection,"
- "Even though the majority of these people did not have any persistent symptoms, their immune system was more sensitive than that of their healthy counterparts."

Hetsouphanh, C., Darley, D.R., Wilson, D.B. et al. Immunological dysfunction persists for 8 months following initial mild-to-moderate SARS-CoV-2 infection. Nat Immunol 23, 210–216 (2022).

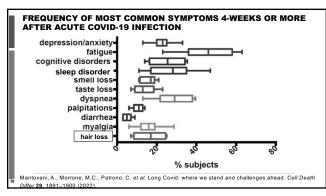






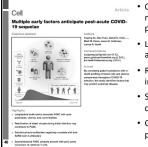








Multiple early factors anticipate post-acute COVID-19 sequelae



- Omics = various disciplines in biology whose names end in omics; e.g., genomics, proteomics, transcriptomics
- Longitudinal multi-omics associate PASC with autoantibodies, viremia, and comorbidities
 Reactivation of latent viruses during initial
- infection may contribute to PASCSubclinical auto-antibodies negatively
- correlate with anti-SARS-CoV-2 antibodies
- Gastrointestinal PASC uniquely present with post-acute expansion of cytotoxic T cells

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OTHER FACTORS THAT MAY CONTRIBUTE TO SYMPTOMS AND TREATMENT OF PASC

- Having COVID-19 can bring to light existing health problems that might not have been noticeable before or worsen the symptoms of a known health condition.
- People who require intensive care for any serious illness, including COVID-19, are more likely to develop weakness, brain and mental health disorders, and other long-term health issues after they leave the hospital.
- Social challenges related to COVID-19, such as isolation and limited access to regular health care, may also underlie some aspects of Long COVID.

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IT'S A QUANDARY

- Unclear why some people develop long-lasting symptoms after having COVID-19, while other people recover completely.
- Understanding the causes of Long COVID is critical for finding ways to prevent, detect, and treat its symptoms.

https://www.nia.nih.gov/health/what-do-we-know-about-long-covid

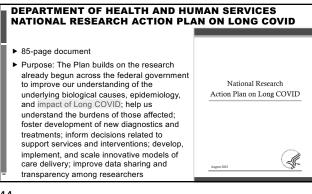
ISSUE FOR LTC RESEARCH AND PASC

► Long Covid associated with more than 200 symptoms and 50 health conditions

- What do we do with this issue in the LTC setting???
- In general, PASC research has excluded older adults, especially the older old (80+ years)
- Also excluded are those with:
 - Multiple complex comorbidities
 - Frailty
 - Disability
 - Dementia
 - Impaired immune function
- All of which are frequent characteristics of LTC residents

Sorensen JM, Crooks VA, Freeman S, Carroll S, Davison KM. A call to action to enhance understanding of long COVID in long-term care home residents. J Am Geriatr Soc. 2022. Jul;70(7):1943-1945.

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Most Common Types of Long Covid Rehab

- Physical therapy
- Pulmonary rehab
- Cognitive rehab
- Mental health support
- Early evidence suggests that some complications, such as heart or lung damage, may improve with time, particularly if patients receive rehabilitative care



REHABILITATION FOR PASC

- Effects of Exercise Rehabilitation in Patients With Long Coronavirus Disease 2019
 European Journal of Preventive Cardiology, May 2022
- Conclusion: Improvements in both cardiorespiratory function and muscular strength were seen in patients with long COVID with a combined aerobic and resistance exercise program. While no control group was compared to the intervention group, this study highlights the potential gains from exercise interventions for this
- Persistent Exerctional Dyspnea and Perceived Exercise Intolerance After Mild COVID-19: A Critical Role for Breathing Dysregulation? Physical Therapy, July 2022
- Conclusion: Breathing dysregulation may help explain COVID-related dyspnea and perceived exercise intolerance after mild infection. The study authors propose that breathing relaxation strategies might be helpful for some individuals, while
 rebreathing therapy may be required for others.

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population.

REHAB STUDIES

- <u>Case Report: Overlap Between Long COVID and Functional Neurological Disorders</u> Frontiers in Neurology, January 2022
- Conclusion: This case report highlights functional neurological disorder as a consideration for patients diagnosed with long COVID. The patient presented with persistent attention and memory difficulties as well as limb dysesthesia after COVID-19 infection. A neurological examination found no organic disorder, and the patient was treated for functional neurological disorder with psychotherapy and physical therapy. The patient's dysesthesia symptoms resolved, and cognition improved.
- Effect of pulmonary rehabilitation approaches on dyspnea, exercise capacity, fatioue, lung functions, and quality of life in patients with covid-19: A systematic review and meta-analysis, Archives of Physical Medicine and Rehabilitation, October 2022
- Conclusion: Pulmonary rehabilitation (exercises, training, education and behavioral changes) were found to be significantly effective in improving dyspnea and exercise capacity in patients with acute and chronic COVID-19 with mild to severe symptoms.
 Fatigue and lung functions were significantly improved in acute COVID-19 patients with mild symptoms.

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PASC REHABILITATION APPROPRIATE CONDITIONS

Lung damage

- Heart damage or inflammation, such as myocarditis or pericarditis
- Cognitive impairments that affect memory or concentration
- Lasting effects from complications:
- Heart attacks,
- Stroke,
- Pulmonary embolism
- Anxiety,
- Depression,
- Muscle/joint pain,
- Chronic fatigue

COVID LONG-HAULERS AND THE SKIN

- Persistent morbidity noted in all systems of the body including skin
 Urticarial and morbilliform eruptions short duration
- ▶ Papulosquamous eruptions, particularly pernio longer-lasting
- American Academy of Dermatology data revealed previously unreported subset of patients who experience long-hauler symptoms in dermatologydominant COVID-19
- Finding raises questions about persistent inflammation; even in patients who initially experienced relatively mild COVID-19
- More studies are needed to understand the long-hauler dermatologic manifestations
- Crift & Bernaber RJ, Land F, Gernelli Against C-P-ACSG. Persistent symptoms in patients after acute COVID-19. JAMA 2020; **326**:603–65. 10. Patimam XD, Carel JA, Weless I, et al. Outcoms of and/ovascular magnetic resonance imaging in patients recently recovered from coronavirus diseas 2091 (COVID-19). AMA Cardor 2020; 5:265–73.

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LESSONS FROM PRACTICE-COVID SKIN MANIFESTATIONS

- Patients with COVID-19 may present with unusual skin manifestations, including urticarial rashes, vesicular lesions, and chilblains on fingers or toes
- These skin and mucous membrane manifestations may be the first sign of COVID-19 disease
- Most cutaneous manifestations of COVID-19 are self-resolving.
- Where treatment is appropriate, medium or high-potency topical corticosteroids, oral antihistamines, or systemic corticosteroids are usually sufficient for symptomatic relief
- Coinciding drug therapy reactions are a possible confounding factor for cutaneous manifestations of COVID-19

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OPEN QUESTIONS REGARDING PASC AND COVID SKIN MANIFESTATIONS

- Occurrence, mechanism, and significance of SARS-CoV-2 persistence in different organs?
- Mechanisms, targets, and significance of autoimmune reactions?
- ► Role of other viruses?
- Impact of host genetics and microbiome?
- Occurrence and severity of PASC after infection with future variants?
- Preventive and therapeutic approaches?
- Years and years of research to come for the regarding PASC

ISSUES WITH RESEARCH IN LTC RESIDENTS

- Research design and interpretation of long COVID outcomes for LTC residents require special consideration of their complex comorbidities and diverse physical, psychological, and social care needs
- Concerns regarding communication impairments that limit self-reporting of symptom
- How do providers and bedside clinical teams tease out long COVID symptoms that may be attributed to pre-existing conditions?

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FEW STUDIES IN THE U.S. FOR LTC RESIDENTS WITH PASC

- Research/studies needed asap to address the lack of guidelines for care and rehabilitation for Long COVID patients and residents in the LTC setting
- ► LTC is a unique care setting
- ► Long COVID needs to be addressed in LTC,
- This issue will probably continue considering the ongoing COVID-19 breakouts by continuously emerging variants
- This and many other immense challenges face the LTC sector related to COVID-19 disease
- We're doing the best we can
- https://www.nia.nih.gov/health/what-do-we-know-about-longcovid#funding



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Thank You!

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REFERENCES - PASC

- Mantovani, A., Morrone, M.C., Patrono, C. et al. Long Covid: where we stand and challenges ahead. Cell Death Differ 29, 1891–1900 (2022).
 Peghin M, Palese A, Venturini M, De Martino. Post-COVID-19 symptoms 6 months after acute infection among hospitalized and non-hospitalized patients. Clinical Microbiology and Infection 27 (2021) 1507-1513.
- Carli A, Bernabei R, Landi F, Gemelli Against C-P-ACSG. Persistent symptoms in patients after acute COVID-19. JAMA 2020; 324: 603–05. McDonald LT. Healing after COVID-19: are survivors at risk for pulmonary fibrosis? Am J Physiol Lung Cell Mol Physiol 320: L257–L265, 2021. ×
- McMahon DE, Gallman AE, Hruza GJ, Rosenbach M, Lipoff JB, Desai SR, et al. Long COVID in the skin: a registry analysis of COVID-19 dermatological duration. The Lancet/infection Vol 21 March 2021 313-314.
- Marcn 2021 313-314. Stefanie Deinhardt-Emmer, Daniel Wittschieber, Juliane Santt, Sandra Kleemann, Stefan Elschner, Karoline Frieda Haupt, Vanessa Vau, Clio Häring, Jörgen Rödel, Andreas Henke, Christina Ehhardt, Michael Bauer, Mike Philipp, Nikolaus Gaßler, Sandor Nietzsche, Bettina Löffler, Gita Mall. Early postmortem mapping of SARS-Co-V2 RNA in patients with COVID-19 and the correlation with tissue damage. *eLile*, 2021; 10 DOI: <u>10.7554/eLile.60361</u>

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REFERENCES - PASC

- Su Y, Yuan D, Chen DG, et al. Multiple early factors anticipate post-acute COVID-19 sequelae. *Cell*. 2022;185(5):881-895.e20. doi:10.1016/j.cell.2022.01.014Peiris S, Mesa H, Aysola A, Manivel J, Toledo J, Borges-Sa M, et al. (2021) Pathological findings in organs and tissues of patients with COVID-19. A systematic review. PLoS ONE 16(4):e0250708. https://doi.org/10.1371/journal.pone.0250708
- CDC.gov. Symptoms of Covid-19.
- Millet C, Narvaneni S, Roman S. Symptoms persist in patients two years after COVID-19 infection: a prospective follow-up study. June 16, 2022. Clin Microbiology and Infection/ European Society of Clinical Microbiology and Infectious Diseases. Accessed 10/28/22
- Whiteson JH, Azola A, Barry JT, et.al Multi-disciplinary collaborative consensus guidance statement on the assessment and treatment of cardiovascular complications in patients with post-acute sequelae of SARS-CoV-2 infection (PASC). PM R. 2022 UI;14(7):855-878. doi: 10.1002/pm;12859. Epub 2022 Jul 13. PMID: 35657351; PMCID: PMC9347705.
- Becker C, Beck K, Zumbrunn S, Memma V, et.al. Long Covid 1 year after hospitalization for covid-19: a prospective bicentric cohort study. Swiss Med Weekly 2021;151:w30091.

SKIN AND MUCOCUTANEOUS MANIFESTATIONS

- Nuno-Gonzalez A, Martin-Carrillo P, Magaletsky K, et al. Prevalence of mucocutaneous manifestations in 666 patients with COVID-19 in a field hospital in Spain: oral and palmoplantar findings. Br J Dermatol. 2021;184(1):184-185. doi:10.1111/bjd.19564 ▶ Mylapalli, H.M. Covid tongue: a new symptom of Covid-19. J. Clin. Pharm. Res., 2021,
- 1(2), 36-38.
- Amorim Dos Santos J, Normando AGC, Carvalho da Silva RL, et al. Oral Manifestations in Patients with COVID-19: A Living Systematic Review. J Dent Res. 2021;100(2):141-154. doi:10.1177/0022034520957289
- Varga Z, Endotheliitis bei COVID-19 [Endotheliitis in COVID-19]. Pathologe. 2020;41(Suppl 2):99-102. Genovese G, Moltrasio C, Berti E, Marzano A, V: Skin Manifestations Associated with COVID-19: Current Knowledge and Future Perspectives. Dermatology 2021;237:1-12. doi: 10.1159/000512932. Accessed 1/26/21
- Iranmanesh B, Khalili M, Amiri R, Zartab H, Aflatoonian M. Oral manifestations of COVID-19 disease: A review article.

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SKIN AND MUCOCUTANEOUS MANIFESTATIONS

- × American Academy of Dermatology Association:
- https://www.aad.org/member/practice/coronavirus/registry 2
- Bataille, V., Visconti, A., Murray, B., Bournot, A., Wolf, J., Ourselin, S., & Falchi, M. (2020). Diagnostic value of skin manifestation of SARS-CoV-2 infection. medRxiv. ×
- Black J and Cuddigan J. Skin manifestations of MCOVP2 interview. Black J and Cuddigan J. Skin manifestations with COVID-19: the purple skin and toes that you are seeing may not be deep tissue pressure injury. WCET® Journal 2020;40(2):18-21 Black J., Cuddigan, J., Capasso, V., Cox, J., Delmore, B., Munoz, N., & Pittman, J. on behalf of the National Pressure Injury Advisory Panel (2020). Unavoidable Pressure Injury during COVID-19 Crisis: A Position Paper from the National Pressure Injury Advisory Panel Auditholo chusure micro gom 5
- Panel. Available at www.npiap.com. 5
- Clinical, Laboratory, and Interferon-Alpha Response Characteristics of Patients With Chilblain-like Lesions During the COVID-19 Pandemic
- Biswal JK , Mohanty SK, Satya, Behera SN , et el. Acute Limb Ischemia: A Catastrophic COVID-19 Sequel Leading to Amputation. https://foamcast.org/2020/04/29/covid-19-cutaneous-manifestations-and-covid-toes/

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SKIN AND MUCOCUTANEOUS MANIFESTATIONS

- Young S, Fernandez AP. Skin manifestations of COVID-19. In COVID-19 Curbside Consultants. Cleveland Clinic. Posted May 7, 2020 https://www.ccjm.org/content/early/2020/05/12/ccjm.87a.ccc03.
- Varga Z. Endotheliitis bei COVID-19 [Endotheliitis in COVID-19]. Pathologe. 2020 Dec;41(Suppl 2):99-102. German. doi: 10.1007/s00292-020-00875-9. PMID: 33306138; PMCID: PMC7731145.
- Dermatology Solutions: Emerging Skin Manifestations of COVID-19.
- https://www.dermsolutionstx.com/covid.
- Marzano AV, Cassano N, Genovese G, Moltrasio C, Vena GA. Cutaneous manifestations in patients with COVID-19: A preliminary review of an emerging issue. Br J Dermatol 2020; published online June 1. DOI:10.1111/bjd.19264.
- . Khezri, Mohammad Rafi; Ghasemnejad-Berenji, Morteza PhD; Jafari, Reza PhD Skin Tissue: A Place for SARS-CoV-2 to Multiply and Transmit?, Advances in Skin & Wound Care: October 2021 Volume 34 Issue 10 p 513-514

SKIN AND MUCOCUTANEOUS MANIFESTATIONS

- Wollina U, Karadağ AS, Rowland-Payne C, Chiriac A, Lotti T. Cutaneous signs in COVID-19 patients: A review [published online ahead of print, 2020 May 10]. Dermatol Ther. 2020. Galván Casas, C., Catala, A. C. H. G, Carretero Hernández, G., Rodriguez-Jiménez, P.,
- Galván Casas, C., Catala, A. C. H. G., Carretero Hernández, G., Ródriguez-Jiménez, P., Fernández-Nieto, D., Rodriguez-Villa Lario, A., & García-Gavin, J. (2020). Classification of the cutaneous manifestations of COVID-19: a rapid prospective nationwide consensus study in Spain with 375 cases. *British Journal of Dermatology*, 183(1), 71-77.
- Casas G et al. Five common skin manifestations of COVID-19 identified. Br J Dermatol 2020; 183:71–77.
- Clinical and Histopathological Features and Potential Pathological Mechanisms of Skin Lesions in COVID-19: Review of the Literature, Dermatopathology,10.3390/dermatopathology7010002, 7.1, (3-16), (2020).
- Feldman SR, Freeman EE. Coronavirus disease 2019 (COVID-19): Cutaneous manifestations and issues related to dermatologic care. <u>https://www.uptodate.com/contents/coronavirus-</u>
- disease-2019-covid-19-cutaneous-manifestations-and-issues-related-to-dermatologic-care
- Giavedoni P, Podlipnik S, Pericàs JM, et al. Skin Manifestations in COVID-19: Prevalence and Relationship with Disease Severity. J Clin Med. 2020;9(10):3261.

61

REFERENCES – OTHER

- Mukerji SS, Solomon IH. What can we learn from brain autopsies in COVID-19?. Neurosci Lett. 2021;742:135528. doi:10.1016/j.neulet.2020.135528.
- Liu J, Li Y, Liu L, et al. Infection of human sweat glands by SARS-CoV-2. Cell Discov 2020;6(1):84.
- COVID-19: Acute limb ischemia <u>https://www.uptodate.com/contents/covid-19-acute-limb-ischemia</u> Accessed 5/22/21
- Dance A. What is a cytokine storm? <u>https://www.knowablemagazine.org/article/health-disease/2020/what-cvtokine-storm</u>
- Oral mucosal lesions in a COVID-19 patient: New signs or secondary manifestations? International Journal of Infectious Diseases, Volume 97, 2020, Pages 326-328.
- Phelan AL, Katz R, Gostin LO. The Novel Coronavirus Originating in Wuhan, China: Challenges for Global Health Governance. JAMA 2020; published online Jan 30. DOI:10.1001/jama.2020.1097.
- Shenoy, N., Luchtel, R. & Gulani, P. Considerations for target oxygen saturation in COVID-19 patients: are we under-shooting?. BMC Med 18, 260 (2020). https://doi.org/10.1186/s12916-020-

01735-2

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REFERENCES – OTHER

- Puntmann VO, Carerj ML, Wieters I, et al. Outcomes of cardiovascular magnetic resonance imaging in patients recently recovered from coronavirus disease 2019 (COVID-19). JAMA Cardiol 2020; 5: 1265–73.
- Gonzalez CE, Gimenez G A, Rodriguez LL, Castro RJG, Martinez A, et al. Acute peripheral arterial thrombosis in COVID-19. Role of endothelial inflammation. Br J Surg. 2020 Sep;107(10):e444e445.
- Arthur JM, Forrest JC, Boehme KW, Kennedy JL, Owens S, Herzog C, et al. (2021) Development of ACE2 autoantibodies after SARSCOV-2 infection. PLoS ONE 16(9): e0257016. https://doi.org/10.1371/journal.open.0257016
- Hirose R, Itoh, Y, Ikegaya H, Miyazaki H, et.al. Differences in environmental stability among SARS-CoV-2 variants of concern: Omicron has higher stability. bioRxiv 2022.01.18.476607; doi: https://doi.org/10.1101/2022.01.18.476607 Accessed 3/11/22
- https://www.cdc.gov/mis/mis-a.html
- Seebacher N, Kirkham J, Smith SD. Lessons from practice-Cutaneous manifestations of COVID-19: diagnosis and management. Med J Aust 2022; 217 (2): 76-78. || doi: 10.5694/mja2.51621
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