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the eyelid dermatitis in parallel with respiratory improvement strongly suggests an association between skin manifestations and COVID-19. However, vitamin supplements (especially vitamin C) can also bring partial benefit in the setting of capillaritis and small-vessel dysfunction. <sup>10</sup>

The present report gives a new insight into COVID-19-associated cutaneous findings and can therefore help clinicians in identifying early signs of the disease. In fact, the great variability of COVID-19-related dermatological disorders gives reason of the difficulties encountered by dermatologists and other physicians in recognizing SARS-CoV2 infection and therefore in treating patients accordingly.

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## **Conflict of interest**

None declared.

## Informed consent

The patients in this manuscript have given written informed consent to publication of their case details.

A. Paganelli,<sup>1,2,\*</sup> [D] F. Garbarino,<sup>1,2</sup> [D] L. Bigi<sup>1</sup>

<sup>1</sup>Clinica Dermatologica, AOU Policlinico di Modena, University of Modena and Reggio Emilia, Modena, Italy, <sup>2</sup>PhD Program in Clinical and Experimental Medicine, University of Modena and Reggio Emilia, Modena,

\*Correspondence: A. Paganelli. E-mail: alessia.paganelli@gmail.com

## References

- 1 Wiersinga WJ, Rhodes A, Cheng AC, Peacock SJ, Prescott HC. Pathophysiology, transmission, diagnosis, and treatment of coronavirus disease 2019 (COVID-19): a review. *JAMA* 2020; 324: 782–793.
- 2 Recalcati S. Cutaneous manifestations in COVID-19: a first perspective. J Eur Acad Dermatol Venereol 2020; 34: e212–e213. https://doi.org/10.1111/ idv.16387
- 3 Freeman EE, McMahon DE, Lipoff JB et al. The spectrum of COVID-19associated dermatologic manifestations: An international registry of 716 patients from 31 countries. J Am Acad Dermatol 2020; 83: 1118–1129.
- 4 Daneshgaran G, Dubin DP, Gould DJ. Cutaneous manifestations of COVID-19: an evidence-based review. Am J Clin Dermatol 2020; 21: 627–639.
- 5 Bouaziz JD, Duong TA, Jachiet M et al. Vascular skin symptoms in COVID-19: a French observational study. J Eur Acad Dermatol Venereol 2020; 34: e451–e452. https://doi.org/10.1111/jdv.16544
- 6 Galván Casas C, Català A, Carretero Hernández G et al. Classification of the cutaneous manifestations of COVID -19: a rapid prospective nationwide consensus study in Spain with 375 cases. Br J Dermatol 2020; 183: 71–77.
- 7 Gianotti R, Veraldi S, Recalcati S et al. Cutaneous clinico-pathological findings in three COVID-19-positive patients observed in the metropolitan area of Milan, Italy. Acta Derm Venerol 2020; 100: adv00124–2.
- 8 Bomhof G, Mutsaers PGNJ, Leebeek FWG et al. COVID-19-associated immune thrombocytopenia. Br J Haematol 2020; 190: e61–e64. https:// doi.org/10.1111/bjh.16850
- 9 Joob B, Wiwanitkit V. COVID-19 can present with a rash and be mistaken for dengue. *J Am Acad Dermatol* 2020; **82**: e177.

10 Schober SM, Peitsch WK, Bonsmann G et al. Early treatment with rutoside and ascorbic acid is highly effective for progressive pigmented purpuric dermatosis. J Dtsch Dermatol Ges 2014; 12: 1112–1119.

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# COVID-19 pandemic and autoimmune bullous diseases: a cross-sectional study of the International Pemphigus and Pemphigoid Foundation

Editor

Autoimmune bullous diseases (AIBDs) are rare and potentially life-threatening chronic inflammatory disorders that are difficult to manage during the COVID-19 outbreak. Our objective was to investigate the associations of outdoor activity restriction, income loss and treatment non-adherence with self-reported outcomes and to determine the satisfaction level with teledermatology platforms in patients with AIBDs during the COVID-19 pandemic.

In this cross-sectional study, English-speaking AIBD patients aged >18 years, who were recruited from the database of the International Pemphigus and Pemphigoid Foundation, were asked to complete a COVID-19 pandemic-related Web-based survey between 30 July 2020 and 1 October 2020. The online poll and its rating system were adapted with minor modifications from Kuang et al.,3 Wang et al.4 and Ruggiero et al.5 Electronic informed consent was obtained from all patients, and the questionnaire was completed anonymously. The study was granted exemption by the Institutional Review Board of the University of Southern California. The primary outcome was deterioration of the disease, determined by the Global Rating of Change. The secondary outcomes included perceived stress and symptoms of anxiety and depression, which were assessed by the visual analogue scale, 2-item Generalized Anxiety Disorder and 2-item Patient Health Questionnaire, respectively. The cut-off points were  $\geq 7$ ,  $\geq 3$  and  $\geq 3$ , respectively, according to previous studies.<sup>3,4,6–8</sup> The tertiary outcome was the satisfaction level of patients using telemedicine platforms (i.e. live interactive video-call visits). Logistic regression was used to estimate associations with adjustments for potential confounders. The effect size is presented as odds ratio, likelihood ratio and 95% confidence interval. P values < 0.05 were considered statistically significant.

Valid questionnaires including location data were collected from a total of 383 patients [276 females and 107 males; aged 19–95 (mean 59.9) years; 207 pemphigus vulgaris, 75 mucous

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Table 1 Associations of outdoor activity restriction, income loss and treatment non-adherence with patient-reported outcomes of AIBDs

Patient-reported outcomes	Unrestric outdoor activity (n = 87)	activity		Partial restriction of outdoor activity $(n = 185)$			Complete restriction of outdoor activity ( <i>n</i> = 111)			LR test	
	n (%)	OR	n (%)	OR (95% CI)	aOR (95% CI)	† n (%)	OR (95% CI)	aOR (95% CI)	† P	аP	
Deteriorated disease	27 (31)	1	65 (35.1)	1.2 (0.7–2.1) P = 0.505	1 (0.6–1.8) P = 0.939	43 (38.7)	1.4 (0.8–2.5) P = 0.261	1 (0.5–1.9) P = 0.932	0.528	0.983	
Perceived stress (VAS, ≥7)	23 (26.4)	1	55 (29.7)	1.2 (0.7–2.1) P = 0.576	1 (0.6–1.8) P = 0.995	46 (41.4)	2 (1.1–3.6) P = 0.028	1.6 (0.8–3) P = 0.183	0.048	0.224	
Anxiety (GAD-2, ≥3)	18 (20.7)	1	69 (37.3)	2.3 (1.3–4.1) P = 0.006	2 (1.1–3.8) P = 0.021	47 (42.3)	2.8 (1.5–5.3) P = 0.001	2.4 (1.2–4.8) P = 0.01	0.003	0.021	
Depression (PHQ-2, ≥3)	18 (20.7)	1	58 (31.4)	1.8 (1–3.2) P = 0.069	1.5 (0.8–2.8) P = 0.198	46 (41.4)	2.7 (1.4–5.2) P = 0.002	1.9 (1–3.8) P = 0.065	0.007	0.175	
Patient-reported outcomes	Income unaffected (n = 254);		Partial income loss (n = 100)			Complete	Complete income loss (n = 29)			it	
	n (%)	OR	n (%)	OR (95% CI)	aOR (95% CI)†	n (%)	OR (95% CI)	aOR (95% CI)	P	a <i>P</i>	
Deteriorated disease	83 (32.7)	1	40 (40)	1.4 (0.9–2.2) P = 0.193	1.4 (0.8–2.4) P = 0.256	12 (41.4)	1.5 (0.7–3.2) P = 0.349	0.3 (0.1–1) P = 0.045	0.336	0.033	
Perceived stress (VAS, ≥7)	71 (28)	1	38 (38)	1.6 (1–2.6) P = 0.066	1.3 (0.7–2.2) P = 0.387	15 (51.7)	2.8 (1.3–6) P = 0.01	1.7 (0.6–4.6) P = 0.275	0.015	0.449	
Anxiety (GAD-2, ≥3)	80 (31.5)	1	40 (40)	1.4 (0.9–2.3) P = 0.129	1.5 (0.9–2.6) P = 0.138	14 (48.3)	2 (0.9–4.4) P = 0.073	1.7 (0.6–4.4) P = 0.296	0.099	0.253	
Depression (PHQ-2, ≥3)	71 (28)	1	42 (42)	1.9 (1.2–3) <i>P</i> = 0.011	2.3 (1.3–4.1) P = 0.003	9 (31)	1.2 (0.5–2.7) P = 0.727	0.9 (0.3–2.5) P = 0.813	0.041	0.009	
Patient-reported outcomes			t Tre		tment non-adhe	rence ( <i>n</i> = 3	nce (n = 30)		LR test		
	n (%	6)	0	R n(%)	) OR	(95% CI)	aOR (95%	CI)† P		a <i>P</i>	
Deteriorated disease	118	(33.4)	1	17 (5	,	(1.2–5.5) 0.013	2 (0.8–4.6) P = 0.122	0.	012	0.123	
Perceived stress (VAS, ≥7)	113 (32)		1	11 (3	,	(0.6–2.7) 0.601	0.7 (0.3–1. P = 0.428	7) 0.	604	0.422	
Anxiety (GAD-2, ≥3)	120 (34)		1 14		•	(0.8–3.6) 0.166	1.8 (0.8–4. P = 0.183	1) 0.	17	0.186	
Depression (PHQ-2, ≥3)	113	(32)	1	9 (30	•	(0.4–2.1) 0.82	0.9 (0.3–2. P = 0.739	2) 0.	32	0.737	

a, adjusted; AIBDs, autoimmune bullous diseases; CI, confidence interval; GAD-2, 2-item Generalized Anxiety Disorder; LR, likelihood ratio; OR, odds ratio; PHQ-2, 2-item Patient Health Questionnaire; VAS, visual analogue scale.

membrane pemphigoid, 59 bullous pemphigoid, 31 pemphigus foliaceus and 11 other AIBDs]. Responses to the online questionnaire came from North America (n=320), Europe (n=26), Asia (n=15), South America (n=11), Africa (n=4), Australia (n=4) and New Zealand (n=3). Eleven (2.8%) patients reported confirmed infection with SARS-CoV-2. 35.2% patients reported a moderate to great exacerbation of their AIBD. The proportions of perceived stress, anxiety and depression were 32.4%, 35% and 31.9%, respectively. Outdoor activity restriction was associated with stress, anxiety and depression. Income loss was associated with stress and

depression. Treatment non-adherence was associated with disease aggravation. After adjustment for confounders, relationships remained between outdoor activity restriction and anxiety as well as between income loss and depression. Confounder adjustment further revealed an association between income loss and disease deterioration (Table 1). Thirty-five (9.1%) patients consistently reported to have consulted a dermatologist through telemedicine platforms, and there was a high satisfaction level (score 7–10) in the majority of patients (72.4–90.3%) who used this virtual healthcare service through videoconferencing (Table 2).

<sup>†</sup>Adjusted for location of study participants, age, gender, education, annual income, marital status, comorbidities, disease type, disease duration, affected body surface area, mucosal involvement, income loss, treatment non-adherence and COVID-19 status. ‡Twelve out of these patients reported increased income

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**Table 2** Six-item questionnaire using a 0–10 scale (score 0–3: negative; 4–6: not bad not good; 7–10: positive): scores reported in 35 autoimmune bullous disease patients

Scores	Α	В	C <sup>1</sup>	$D^2$	E <sup>3</sup>	F
	No. of patients					
0	1	0	2	0	1	1
	(2.9%)	(0%)	(6.9%)	(0%)	(3.2%)	(2.9%)
	4	4	5	1	3	5
	(11.4%) <sup>4</sup>	(11.4%) <sup>4</sup>	(17.2%) <sup>4</sup>	(3.1%) <sup>4</sup>	$(9.7\%)^4$	(14.3%) <sup>4</sup>
1	1	1	1	0	1	0
	(2.9%)	(2.9%)	(3.4%)	(0%)	(3.2%)	(0%)
2	1	3	1	0	1	2
	(2.9%)	(8.6%)	(3.4%)	(0%)	(3.2%)	(5.7%)
3	1	0	1	1	0	2
	(2.9%)	(0%)	(3.4%)	(3.1%)	(0%)	(5.7%)
4	0	3	0	1	0	0
	(0%)	(8.6%)	(0%)	(3.1%)	(0%)	(0%)
	4	4	3	6	0	2
	(11.4%) <sup>5</sup>	(11.4%) <sup>5</sup>	(10.3%) <sup>5</sup>	(18.8%) <sup>5</sup>	(0%) <sup>5</sup>	$(5.7\%)^5$
5	2	1	2	5	0	1
	(5.7%)	(2.9%)	(6.9%)	(15.6%)	(0%)	(2.9%)
6	2	0	1	0	0	1
	(5.7%)	(0%)	(3.4%)	(0%)	(0%)	(2.9%)
7	1	3	1	1	0	4
	(2.9%)	(8.6%)	(3.4%)	(3.1%)	(0%)	(11.4%)
	27	27	21	25	28	28
•	(77.1%) <sup>6</sup>	(77.1%) <sup>6</sup>	(72.4%) <sup>6</sup>	(78.1%) <sup>6</sup>	(90.3%) <sup>6</sup>	(80%) <sup>6</sup>
8	4	2	2	-		•
	(11.4%)	(5.7%)	(6.9%)	(18.8%)	(6.5%)	(2.9%)
9	4	5	4	0	3	0
10	(11.4%)	(14.3%)	(13.8%)	(0%)	(9.7%)	(0%)
10	18	17	14	18	23	23
	(51.4%)	(48.6%)	(48.3%)	(56.2%)	(74.2%)	(65.7%)

(A) I was satisfied with the attention paid by the doctor to my disease. (B) I was satisfied with the time spent by the doctor with me. (C) I was satisfied with the treatment I received. (D) I was satisfied with the convenience compared to a regular clinic visit. (E) I was satisfied with the coronavirus safety compared to a regular clinic visit. (F) What is the likelihood that you would use this dermatology telemedicine service again?

Our results emphasize the negative impact of the COVID-19 outbreak on health outcomes in patients with AIBDs and indicate a high satisfaction with telemedicine platforms at this difficult time, although the prevalence rates of stress, anxiety and depression were comparable to the general population during the COVID-19 pandemic (i.e. 29.6%, 31.9% and 33.7%, respectively). Appropriate healthcare delivery solutions, including a public mental health response, are required to improve the health of vulnerable individuals in the COVID-19 era.

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M. Kasperkiewicz,<sup>1,\*</sup> M. Yale,<sup>2</sup> R. Strong,<sup>2</sup> D. Zillikens,<sup>3</sup> D.T. Woodley,<sup>1</sup> A. Recke<sup>3</sup>

<sup>1</sup>Department of Dermatology, Keck School of Medicine, University of Southern California, Los Angeles, CA, USA, <sup>2</sup>International Pemphigus and Pemphigoid Foundation, Sacramento, CA, USA, <sup>3</sup>Department of Dermatology, University of Lübeck, Lübeck, Germany \*Correspondence: M. Kasperkiewicz. E-mail: michael.kasperkiewicz@

med.usc.edu

# References

- 1 Kasperkiewicz M, Schmidt E, Fairley JA et al. Expert recommendations for the management of autoimmune bullous diseases during the COVID-19 pandemic. J Eur Acad Dermatol Venereol 2020; 34: e302–e303.
- 2 Kasperkiewicz M. COVID-19 outbreak and autoimmune bullous diseases: a systematic review of published cases. J Am Acad Dermatol 2021; 84: 563–568.

<sup>&</sup>lt;sup>1</sup>Six patients did not provide a score. <sup>2</sup>Three patients did not provide a score. <sup>3</sup>Four patients did not provide a score. <sup>4-6</sup>Numbers (in italic) represent the sum of scores 0–3, 4–6 and 7–10, respectively.

Letters to the Editor e421

- 3 Kuang Y, Shen M, Wang Q *et al.* Association of outdoor activity restriction and income loss with patient-reported outcomes of psoriasis during the COVID-19 pandemic: a web-based survey. *J Am Acad Dermatol* 2020; **83**: 670–672.
- 4 Wang Q, Luo Y, Lv C et al. Nonadherence to treatment and patient-reported outcomes of psoriasis during the COVID-19 epidemic: a web-based survey. Patient Prefer Adherence 2020; 14: 1403–1409.
- 5 Ruggiero A, Megna M, Annunziata MC et al. Teledermatology for acne during COVID-19: high patients' satisfaction in spite of the emergency. J Eur Acad Dermatol Venereol 2020; 34: e662–e663.
- 6 Lesage FX, Berjot S. Validity of occupational stress assessment using a visual analogue scale. *Occup Med (Lond)* 2011; **61**: 434–436.
- 7 Plummer F, Manea L, Trepel D, McMillan D. Screening for anxiety disorders with the GAD-7 and GAD-2: a systematic review and diagnostic metaanalysis. Gen Hosp Psychiatry 2016; 39: 24–31.
- 8 Kroenke K, Spitzer RL, Williams JBW. The patient health questionnaire-2: validity of a two-item depression screener. Med Care 2003; 41: 1284–1292.
- 9 Salari N, Hosseinian-Far A, Jalali R et al. Prevalence of stress, anxiety, depression among the general population during the COVID-19 pandemic: a systematic review and meta-analysis. Global Health 2020; 16: 57.

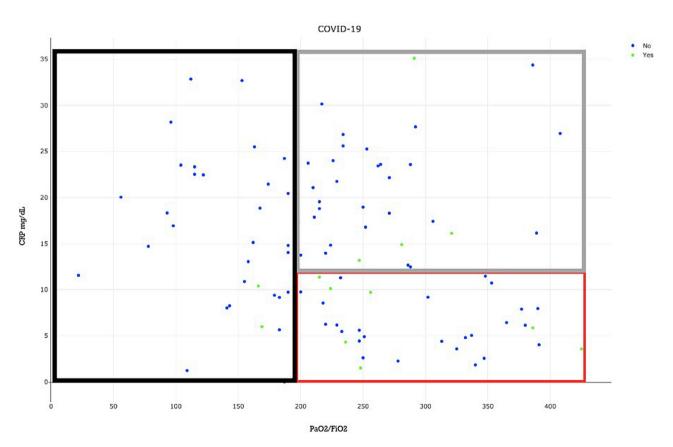
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# Follow-up of dermatological manifestations in non-critical hospitalized patients with COVID-19 pneumonia and their prognostic correlation with disease severity

Dear Editor,

COVID-19 is currently one of the main causes of death worldwide. This virus affects mainly the lower respiratory system, but significant damage to other organs has been observed. Cutaneous manifestations related to the aforementioned viral infection have been reported with an incidence that ranges between 0.20% and 20%. The period between the appearance of cutaneous lesions and COVID-19 infection remains uncertain. 1,2

With the information that exists, one can speculate that cutaneous manifestations of COVID-19 can be classified into two groups



**Figure 1** In this graphic, patients with cutaneous manifestations (blue dots) and without cutaneous manifestations (green dots) are plotted according to their prognosis using clinical and biochemical variables. The red box represents patients with a P/F ratio >200 and CRP <11 mg/dL. The grey box represents patients with a P/F ratio >200 and CRP >11 mg/dL. The black box represents patients with a P/F ratio <200 and CRP >11 mg/dL.