

Figure 1 Clinical appearance of an asymptomatic patient with atopic dermatitis before starting dupilumab, showing no abnormalities (a). Appearance of the same eye after Lissamine Green conjunctival staining, highlighting the presence of moderate ocular surface alterations (white arrow) on the marginal portion of the lower tarsal conjunctiva (lid wiper epitheliopathy) prior to dupilumab administration (b). The same eye after 16 weeks of treatment showing severe epitheliopathy of the lower tarsal conjunctiva (yellow arrow) in addition to the previous alterations: DIOSD (c).

with AD suitable for systemic therapy with dupilumab. A correct recognition of early ocular alterations can direct us towards a targeted treatment, thus obtaining a good control of both the symptoms and the ocular clinical picture to avoid the withdrawal of such an effective drug.

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References

- 1 Worm M, Simpson EL, Thaçi D et al. Efficacy and safety of multiple dupilumab dose regimens after initial successful treatment in patients with atopic dermatitis: a randomized clinical trial. JAMA Dermatol 2020; 156:131–43.
- 2 Akinlade B, Guttman-Yassky E, de Bruin-Weller M et al. Conjunctivitis in dupilumab clinical trials. Br J Dermatol 2019; 181:459–73
- 3 Zirwas MJ, Wulff K, Beckman K. Lifitegrast add-on treatment for dupilumab-induced ocular surface disease (DIOSD): a novel case report. JAAD Case Rep 2018; 5:34-6.

- 4 Maudinet A, Law-Koune S, Duretz C et al. Ocular surface diseases induced by dupilumab in severe atopic dermatitis. Ophthalmol Ther 2019; 8:485–90.
- 5 Miller KL, Walt JG, Mink DR et al. Minimal clinically important difference for the ocular surface disease index. Arch Ophthalmol 2010; 128:94—101.
- 6 Chen JJ, Applebaum DS, Sun GS, Pflugfelder SC. Atopic keratoconjunctivitis: a review. J Am Acad Dermatol 2014; 70:569–75.
- 7 Raffi J, Suresh R, Fishman H et al. Investigating the role of allergic contact dermatitis in residual ocular surface disease on dupilumab (ROSDD). Int J Womens Dermatol 2019; 5:308–13.
- 8 Hu Y, Matsumoto Y, Adan ES et al. Corneal in vivo confocal scanning laser microscopy in patients with atopic keratoconjunctivitis. Ophthalmology 2008; 115:2004–12.

Funding: None

Conflicts of interest: we declare that we have no conflicts of interest.

'Not relevant' responses in the era of COVID-19: are we underestimating Dermatology Life Quality Index values?

DOI: 10.1111/bjd.20705

DEAR EDITOR, Since March 2020, the UK and Ireland have entered a series of lockdowns in the wake of the coronavirus pandemic with restrictions resulting in the closure of nonessential retail, hospitality and sports with employees encouraged to work from home. 1,2 Patients on biologic and systemic immunosuppression often restricted their lifestyles more than the general public. 3 The Dermatology Life Quality

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Index (DLQI) is a measure of health-related quality of life used to guide treatment decisions and is an important component of data collected for clinical trials and registries. Previous studies have demonstrated that 'not relevant' responses (NRRs) in the DLQI can impact the validity of treatment decisions. Given the restrictions imposed on our patients during this time, we hypothesized that the number of NRRs in the DLQI increased in the era of COVID-19 with the potential to affect treatments offered and analysis of data from clinical trials and registries.

Our primary aim was to identify the number of NRRs prior to and during COVID-19 restrictions and our secondary aim was to assess the difference between the DLQI and the DLQI-R (DLQI-Relevant). We completed a retrospective chart review for patients with stable plaque psoriasis attending the specialist psoriasis clinic in our centre. We defined stable disease as a change in Psoriasis Area and Severity Index (PASI) \leq 4, which was demonstrated previously to have no effect or a small effect on DLQI values.^{6,7} The patients' most recent DLQI score prior to restrictions was identified and compared with their most up-to-date DLQI score during lockdown. We applied the previously reported modified DLQI-R scoring system to calculate a value taking into account the NRRs.8 The DLQI-R assumes a measure of positivity in each NRR with multiplication by a certain factor depending on the number of NRRs. Statistical analysis was carried out using SPSS Statistics for Macintosh V27.0 (IBM Corp., Armonk, NY, USA). Student's paired t-test was used to compare continuous data and Chi-square to compare frequencies in the NRRs. As this study involved the retrospective analysis of routine clinical data, ethical approval was not required.

We identified 68 patients with psoriasis in four weekly specialist clinics with completed DLQIs before COVID-19 and during restrictions. Sixteen patients were excluded from analysis due to a change in PASI $> 4\cdot0$. In the remaining 52 patients, there were 28 females and 24 males with an average age of 55·29 (range 27–82) years. There were 40 patients on a biologic, 10 patients on a systemic agent and two patients using topical therapies. The mean interval between DLQIs was 19·83 (range 13–26) months.

The mean PASI score prior to COVID-19 was $2\cdot16$ (range $0-7\cdot9$, SD $1\cdot77$; Table 1). The mean DLQI score prior to COVID-19 was $3\cdot13$ (range 0-14, SD $3\cdot77$) with the mean NRRs $0\cdot62$ (range 0-3, SD $1\cdot05$). Application of the DLQI-R increased the mean to $3\cdot43$ (range $0-16\cdot25$, SD $4\cdot11$) giving a mean change with the DLQI-R of $0\cdot29$ (range $0-3\cdot25$, SD $0\cdot69$). There were NRRs in the sport (n = 9, 17%), working/studying (n = 7, 13%), sexual difficulties (n = 7, 13%), partner/friends/relatives (n = 6, 12%), treatment (n = 1, 2%), clothes (n = 1, 2%) and shopping/home/garden (n = 1, 2%) questions. There were no NRRs for the social/leisure question.

The mean PASI score during restrictions was 2.09 (range 0-7.8, SD 2.12) (Table 1). The mean DLQI score during restrictions was 3 (range 0-14, SD 3.99) with the mean NRRs 1.27 (range 0-8, SD 2.06). Application of the DLQI-R increased the mean to 3.63 (range 0-17.5, SD 4.71) giving a mean change with the DLQI-R of 0.63 (range 0-12, SD

Table 1 Mean PASI, DLQI, number of NRRs, DLQI-R and change pre-COVID-19 and during lockdown restrictions

Mean DLQI, NRRs, DLQI-R and change pre-COVID-19 and

during restrictions			
	Mean	Range (SD)	P-value ^a
PASI pre-COVID	2.16	0-7.9 (1.77)	0.745
PASI during restrictions	2.09	0-7.8 (2.12)	
DLQI pre-COVID	3.13	0-14 (3.77)	< 0.001
DLQI during restrictions	3	0-14 (3.99)	
DLQI change	-0.13	-11-14 (3.36)	
NRRs pre-COVID	0.62	0-3 (1.05)	
NRRs during restrictions	1.27	0-8 (2.06)	
DLQI-R pre-COVID	3.43	0-16-25 (4-11)	
DLQI-R during restrictions	3.63	0-17.5 (4.71)	
DLQI-R change pre-COVID	0.29	0-3.25 (0.69)	
DLQI-R change	0.63	0-12 (1.88)	
during restrictions			

NRRs pre-COVID-19 and during restrictions, n (%)				
	Pre- COVID	During restrictions	P-value ^b	
Item 3: Going shopping/looking after home/garden	1 (2)	6 (12)	< 0.001	
Item 4: Clothes	1 (2)	5 (10)	< 0.001	
Item 5: Social/leisure activities	0 (0)	7 (13)	< 0.001	
Item 6: Sport	9 (17)	16 (31)	< 0.001	
Item 7: Working/studying	7 (13)	11 (21)	0.003	
Item 8: Problems with partner/close friends/relatives	6 (12)	8 (15)	0.001	
Item 9: Sexual difficulties	7 (13)	11 (21)	0.003	
Item 10: Treatment	1 (2)	2 (4)	< 0.001	

DLQI, Dermatology Life Quality Index; DLQI-R, DLQI-Relevant; NRRs, 'not relevant' responses; PASI, Psoriasis Area and Severity Index. ^aCalculated using Student's paired t-test; ^bcalculated using Chi-square.

1.88). NRRs were seen in all questions including sport (n = 16, 31%), working/studying (n = 11, 21%), sexual difficulties (n = 11, 21%), partner/friends/relatives (n = 8, 15%), social/leisure (n = 7, 13%), shopping/garden/home (n = 6, 12%), clothes (n = 5, 10%) and treatment (n = 2, 4%).

Using Student's paired t-test for analysis, DLQI values during restrictions were significantly lower than prior to COVID-19 (P < 0.001) (Table 1). There was a statistically significant increase in the number of NRRs during restrictions (P < 0.001) with a corresponding significant increase in DLQI-R (P < 0.001) and in change between DLQI-R and DLQI (P < 0.001). The change in DLQI-R is likely to be underestimated due to the number of 0 DLQI responses pre-COVID-19 (P = 16) and during restrictions (P = 16). Prior to coronavirus there were no NRRs for the social/leisure question but this increased to seven during restrictions. The number of NRRs increased in every category during restrictions.

This is a small study with limitations including the number of patients with a DLQI of 0. However, our study has demonstrated a significant decrease in DLQI values, increase in the

NRRs in the DLQI and change between DLQI and DLQI-R during lockdown restrictions. Given the curtailment in nonessential retail, hospitality, gyms and with large numbers of patients working from home it is important for clinicians to identify the number of NRRs in the DLQI and acknowledge the potential effect on treatment decisions and data collection for disease registries and clinical trials during the pandemic.

Acknowledgments: Open access funding provided by IReL.

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References

- 1 Department of the Taoiseach, gov.ie. Ireland placed on full Level 5 restrictions of the Plan for Living with COVID-19. 30 December 2020 [online]. Available at: https://www.gov.ie/en/press-release/066ce-ireland-placed-on-full-level-5-restrictions-of-the-plan-for-living-with-covid-19/ (accessed 10 March 2021).
- 2 Prime Minister's Office, gov.uk. Prime Minister's statement on coronavirus (COVID-19): 19 December 2020 [online]. Available at: https://www.gov.uk/government/speeches/prime-ministersstatement-on-coronavirus-covid-19-19-december-2020 (accessed 10 March 2021).
- 3 Mahil SK, Yates M, PsoProtect LSM et al.; PsoProtect, CORE-UK study groups. Risk-mitigating behaviours in people with inflammatory skin and joint disease during the COVID-19 pandemic differ by treatment type: a cross-sectional patient survey. Br J Dermotol 2021; 185:80–90.
- 4 Smith C, Yiu Z, Bale T et al. British Association of Dermatologists guidelines for biologic therapy for psoriasis 2020: a rapid update. Br J Dermatol 2020; 183:628–37.
- 5 Rencz F, Poór A, Péntek M et al. A detailed analysis of 'not relevant' responses on the DLQI in psoriasis: potential biases in treatment decisions. J Eur Acad Dermatol Venereol 2018; 32:783–90.
- 6 Mahil SK, Wilson N, Dand N et al.; BADBIR study group and the PSORT consortium. Psoriasis treat to target: defining outcomes in psoriasis using data from a real-world, population-based cohort study (the British Association of Dermatologists Biologics and Immunomodulators Register, BADBIR). Br J Dermatol 2019; 182:1158-66.
- 7 Mattei P, Corey K, Kimball A. Psoriasis Area Severity Index (PASI) and the Dermatology Life Quality Index (DLQI): the correlation between disease severity and psychological burden in patients treated with biological therapies. J Eur Acad Dermatol Venerol 2013; 28:333–7.
- 8 Rencz F, Gulácsi L, Péntek M et al. Proposal of a new scoring formula for the Dermatology Life Quality Index in psoriasis. Br J Dermatol 2018; 179:1102–8.

Funding sources: N.K. is in receipt of grant funding from the City of Dublin Skin and Cancer Hospital Charity.

Conflicts of interest: none to declare.

Data availability: the data that support the findings of this study are available from the corresponding author upon reasonable request.

A national audit of oral propranolol for the treatment of infantile haemangiomas

DOI: 10.1111/bjd.20738

Dear Editor, Infantile haemangiomas (IH) affect up to 4% of infants and are the most common tumour of infancy. Although self-limiting, some lesions can lead to visual impairment, airway obstruction, ulceration or cosmetic disfigurement, and require intervention. Oral propranolol – a nonselective beta blocker – was serendipitously identified in 2008 as an effective treatment for IH and is now recommended as first-line therapy for complicated lesions. ^{2,3}

Despite the success of therapy, there has been trepidation among dermatologists and paediatricians about how to initiate oral propranolol in the neonate population. Accordingly, in 2018 the British Society of Paediatric Dermatology (BSPD) issued unified consensus guidelines for prescribing propranolol for the treatment of IH. We undertook a national audit to determine current prescribing patterns 2 years after the publication of this guidance.

The aims of the audit were to ascertain indications for the initiation, baseline investigations, dosing and daycare admission rates for the induction of oral propranolol. The audit was accomplished in December 2020 using Survey Monkey. Six clinical scenarios with associated questions were included in the online questionnaire. Sixty-five dermatologists (227 members of the BSPD and 103 Irish Association of Dermatology members) completed the survey.

The first case described an IH of the eyelid obstructing the field of vision (Figure 1a). Ninety-five per cent of respondents agreed that oral propranolol was indicated; 3% considered topical timolol to be the first-line therapy.

The second case was an uncomplicated lesion on the abdomen (Figure 1b). Eighty-nine per cent of respondents would offer the parents reassurance, 4% would consider oral propranolol and 6% would propose treatment with topical timolol.

The third case illustrated an ulcerated lesion on the trunk (Figure 1c). In this case, physicians could choose from multiple applicable answers. Ninety-four per cent agreed that oral propranolol was indicated; 84% would also recommend topical (36%) or oral (33%) antibiotics, and topical corticosteroids (25%). Eight per cent would consider using oral corticosteroids, 5% topical timolol and 6% pulsed dye laser.

The fourth image displayed a nasal tip lesion in a healthy 6-week-old infant (Figure 1d). Ninety-three per cent of respondents agreed that oral propranolol was indicated. Prior to starting propranolol, responders were offered a range of