

Improved treatment adherence and allergic disease control during a COVID-19 pandemic lockdown

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To the Editor:

Modern hectic lifestyles have created challenges for parents of allergic children, as treatment adherence and optimal disease control are closely tied to adequate caregiver supervision. Environmental allergens exposure in schools and outdoor environments are also difficult to control but impacts disease outcomes.¹ Improvements in disease symptoms are often reported during school vacations,² when children have more time at home under a caregiver's supervision. This study aimed to evaluate the impact of an 8-week (April to June 2020) COVID-19-induced nation-wide lockdown in Singapore on allergic disease control in children. Universal masking and movement restrictions (closure of schools, workplaces, retail and entertainment facilities) were implemented. This presented a unique opportunity to evaluate the impact of increased caregiver supervision, school closures, reduction of outdoor exposures and increased indoor time on allergic diseases control.

An anonymous self-administered questionnaire, hosted on a secure online portal (FormSG), was disseminated to the general public through several media platforms in the one month after nationwide lockdown measures were lifted. Parents of 0-to-18-year-old children, or adolescents themselves, with allergic disorders (Eczema, Asthma and Allergic Rhinitis (AR)) were invited to complete the questionnaire anonymously. Survey completion implied consent to participate. The study received ethics approval by the NHG Domain Specific Review Board, Singapore (Reference number 2020/00717).

Data on demographics, caregiving arrangements, self-reported symptoms of allergic diseases and self-perceived reasons behind the changes in disease status were collected. Perceived treatment adherence was measured using a 10-point rating scale evaluating adherence before and during lockdown. Disease control was assessed through validated symptom scoring tools – the Patient-Oriented Eczema Measure (POEM) for eczema,³ Asthma Control Test (ACT) for asthma⁴ and Total Nasal Symptom Score (TNSS) for AR.⁵

Data was analyzed using SPSS Version 26.0 (IBM Corp, New York, NY, USA). Pearson's chi-square test was used for categorical variables and Wilcoxon Sign Rank Test for symptom score comparisons. Differences in medians between groups reporting different disease outcomes before and during lockdown were analysed with the Kruskal Wallis test.

A total of 173 parents/adolescents: 89 (51.4%) males and 84 (48.6%) females aged 2 months to 18 years participated in this survey (Supplementary Table 1). The majority of subjects (93.1%) received their treatment under adult supervision.

A total of 41.7% of subjects with eczema reported symptomatic improvement (Figure 1A) and improved treatment adherence during the lockdown [median scores 7 (IQR: 5-8) and 8 (IQR: 7-9) before and during lockdown respectively; $p < 0.001$]. This was significantly associated with a better overall perception of disease control ($p < 0.001$) and improved POEM scores: 9 (IQR: 3-13) to 6 (IQR: 3-10) ($p = 0.001$) (Figure 1B). Significant improvement in itch ($p = 0.005$), bleeding ($p = 0.002$), cracked skin ($p = 0.017$) and flaky skin ($p = 0.03$) were noted, but frequency of topical steroid use was not significantly reduced ($p = 0.217$) (Supplementary Table 2).

Subjects who reported perceived improvement in eczema control had a significantly higher POEM scores before lockdown (median 13, IQR 7 - 18) compared with those who reported worsening (median 5, IQR 1 - 8) or no change in overall eczema control (median 8, IQR 3 - 11) ($p < 0.001$) (Table 1).

Reduced exposure to triggers such as heat/sweat and physical activity (93.8%) and more time for skin care treatment (72.9%) were the most commonly cited reasons for improvement, while increased exposure to indoor triggers such as dust, and heat/sweat were the most commonly cited reasons for deterioration (Figure 2).

Asthmatic patients reported an overall significant improvement in perceived treatment adherence during the lockdown [median scores 9 (IQR: 8-10) and 10 (IQR: 9-10) respectively; $p = 0.011$] (Supplementary Figure 1) but no improvement in overall asthma control ($p = 0.12$) or ACT scores ($p = 0.063$) (Figure 1B). Patients with poorer asthma control before lockdown were more likely to benefit from lockdown – baseline median ACT score was 17 (IQR: 15 - 21) in those who improved, compared to subjects without

improvement (median 24) ($p = 0.001$) (Table 1). Reduced exposure to infections, better sleep, more time for treatment administration, better diet and more time to seek medical attention were the main reasons for asthma improvement, while deterioration was solely attributed to increased indoor dust exposure.

There was no difference in overall median treatment adherence scores in subjects with AR ($p = 0.201$), but better adherence was associated with improved overall perception of disease control ($p = 0.044$). There were no differences in median TNSS scores: [3 (IQR: 2-4 and IQR: 2-5 before and during lockdown respectively) ($p = 0.299$)] (Figure 1B), or when stratified by baseline disease status ($p = 0.149$) (Table 1). All subjects reported reduced indoor dust exposure, better treatment adherence, improved diet and sleep patterns as main reasons for improved AR control. Symptom deterioration was attributed to increased indoor dust exposure, avoidance of medical consultation and lack of caregiver supervision due to remote working demands. Lack of access to medical care was not cited as a reason for poor disease control in any of the allergic disorders.

The COVID-19 lockdown had an unexpected positive impact on treatment adherence and disease symptoms in children with eczema and asthma. Eczema treatment regimens are often complex and time-consuming,⁶ and non-adherence to treatment is a major reason for treatment failure.⁷ The mandatory home confinement serendipitously afforded flexibility of time and increased caregiver supervision, which likely translated to improved treatment adherence and better eczema control, particularly in those with moderately severe eczema (POEM score 8-16³). Reduction of outdoor heat and UVR exposure, humidity, perspiration and exercise, which are known eczema triggers, likely contributed to this improvement as well.

Poorly controlled asthmatics appear to benefit more from reduced exposure to viral respiratory infections, attributable to social distancing, enhanced mask wearing and hand hygiene practices during this lockdown period. Other studies have also observed a reduction in viral infections, Emergency Department visits and admissions for wheezing and asthma exacerbations during the COVID-19 pandemic.^{8, 9}

Reduced healthcare access or medication supply disruptions were not reasons for worsening disease control. This was likely mitigated by enhanced telemedicine practices which is particularly suitable for stable allergic diseases,¹⁰ and home delivery of medications in the local setting which ensured continued access to medical care.

This was a small pilot cross-sectional study without a longitudinal component, and hence may not be sufficiently powered and does not measure sustainability over time. The use of an anonymous self-administered questionnaire might also introduce recall and selection bias. Data on socio-economic status and objective measures of disease, were intentionally omitted from the survey design to minimize subject burden. Furthermore, the study was designed to measure self-perception of treatment adherence and disease symptoms, which are closely correlated with quality of life. Survey respondents may also have been self-selected for better disease control at baseline, thus limiting our ability to measure a tangible improvement in asthma and AR scores.

However, this study has generated important insights into the benefits of extended home-based care on allergic disease control; and the way the COVID-19 pandemic has shaped patient behaviour and its impact on self-perceived allergic disease control, which may in turn influence disease management strategies in the longer term. Extended home-based care can be facilitated by flexible remote school and work arrangements, which are now gaining acceptance worldwide. Larger epidemiological studies are needed to determine if such arrangements lead to reduced healthcare costs, hospital admissions, quality of life and increased work productivity and its associated economic benefits.

Keywords : COVID-19, Allergic Disease, Quarantine

Signature:

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References

1. Muraro A, Clark, A., Beyer, K., Borrego, L.M., Borres, M., Lødrup Carlsen, K.C., Carrer, P., Mazon, A., Rancè, F., Valovirta, E., Wickman, M., Zanchetti, M. The management of the allergic child at school: EAACI/GA2LEN Task Force on the allergic child at school: EAACI/GA2LEN Task Force on the allergic child at school. *Allergy* 2010; 65: 9.
2. Julious SA, Campbell MJ, Bianchi SM, et al. Seasonality of medical contacts in school-aged children with asthma: association with school holidays. *Public Health* 2011; 125: 769-776. 2011/11/01. DOI: 10.1016/j.puhe.2011.08.005.
3. Charman CR, Venn AJ and Williams HC. The patient-oriented eczema measure: development and initial validation of a new tool for measuring atopic eczema severity from the patients' perspective. *Arch Dermatol* 2004; 140: 1513-1519. 2004/12/22. DOI: 10.1001/archderm.140.12.1513.
4. Liu AH, Zeiger R, Sorkness C, et al. Development and cross-sectional validation of the Childhood Asthma Control Test. *J Allergy Clin Immunol* 2007; 119: 817-825. 2007/03/14. DOI: 10.1016/j.jaci.2006.12.662.
5. Pfaar O, Demoly P, Gerth Van Wijk R, et al. Recommendations for the standardization of clinical outcomes used in allergen immunotherapy trials for allergic rhinoconjunctivitis: an EAACI Position Paper. *Allergy* 2014; 69: 854-867. DOI: 10.1111/all.12383.
6. Santer M, Burgess H, Yardley L, et al. Managing childhood eczema: qualitative study exploring carers' experiences of barriers and facilitators to treatment adherence. *J Adv Nurs* 2013; 69: 2493-2501. 2013/03/27. DOI: 10.1111/jan.12133.
7. Krejci-Manwaring J, Tusa MG, Carroll C, et al. Stealth monitoring of adherence to topical medication: adherence is very poor in children with atopic dermatitis. *J Am Acad Dermatol* 2007; 56: 211-216. 2007/01/17. DOI: 10.1016/j.jaad.2006.05.073.
8. Abe K, Miyawaki A, Nakamura M, et al. Trends in hospitalizations for asthma during the COVID-19 outbreak in Japan. *The Journal of Allergy and Clinical Immunology: In Practice* 2020. DOI: 10.1016/j.jaip.2020.09.060.
9. Taquechel K, Diwadkar AR, Sayed S, et al. Pediatric Asthma Health Care Utilization, Viral Testing, and Air Pollution Changes During the COVID-19 Pandemic. *The Journal of Allergy and Clinical Immunology: In Practice* 2020. DOI: 10.1016/j.jaip.2020.07.057.
10. Hare N, Bansal P, Bajowala SS, et al. Work Group Report: COVID-19: Unmasking Telemedicine. *J Allergy Clin Immunol Pract* 2020; 8: 2461-2473 e2463. 2020/07/01. DOI: 10.1016/j.jaip.2020.06.038.

Table 1. Patient rated scores before and during lockdown and overall perceived disease status during lockdown

Score (Dis- ease)	Subjects	Improved disease status during lock- down	Improved disease status during lock- down	Worsened disease status during lock- down	Worsened disease status during lock- down	Stayed the same during lock- down	Stayed the same during lock- down	p- value*
		Median score (IQR)	n	Median score (IQR)	n	Median score (IQR)	n	
POEM Score Before lockdown	POEM Score 115	POEM Score 13 (7 – 18)	POEM Score 48	POEM Score 5 (1 – 8)	POEM Score 26	POEM Score 8 (3 – 11)	POEM Score 41	POEM Score < 0.00
During lockdown		6 (2 - 8)		11 (6 – 15)		7 (2 – 10)		
ACT Score Before lockdown	ACT Score 29	ACT Score 17 (15 – 21)	ACT Score 11	ACT Score 24	ACT Score 3	ACT Score 24 (22-25)	ACT Score 15	ACT Score 0.001
During lockdown		23 (20 - 25)		24		25 (23 - 25)		
TNSS Before lockdown	TNSS 68	TNSS 4 (2 – 5)	TNSS 21	TNSS 3 (2 – 4)	TNSS 17	TNSS 2 (2 – 5)	TNSS 30	TNSS 0.149
During lockdown		2 (2 – 3)		5 (5 – 7)		2.5 (2 – 5)		

AD: Atopic Dermatitis

POEM: Patient Oriented Eczema Measure

ACT: Asthma Control Test

TNSS: Total Nasal Symptom Score

Bold p values indicate statistical significance ($p < 0.05$)

**Analysed by Kruskal-Wallis Test*

Figure 1. Control of allergic disease before and during lockdown.

Figure legend.

Figure 1A shows the breakdown of perceived overall changes in allergic disease control before and during the lockdown period by individual allergic disorders. Data are presented by percentages of the total number of individuals with a particular allergic disorder.

Figure 1B show comparisons of the validated symptom scores: Patient-Oriented Eczema Measure (POEM) scores for eczema, Asthma Control Test (ACT) scores for asthma and Total Nasal Symptom Score (TNSS) for allergic rhinitis before and during the lockdown period. POEM scores range from 0 – 28, with a higher score indicating poorer disease control. ACT

scores range from 5 – 25 with higher scores indicating better disease control. TNSS scores range from 0 – 12, with higher scores indicating poorer disease control.

Figure 2. Factors implicated in Eczema (AD) control

Figure legend.

More time for administration of skin treatment (n=35) and reduced exposure to triggers (n=45) (4A) such as physical activity (n=22) and heat/sweating (n=35) (4B) were the main reasons for improved eczema control during the lockdown period. Increased exposure to indoor triggers (4C) such as house dust mites (HDM) (n=16) and heat/sweat (n=15) (4D) were the main reasons for poor eczema control. Other reported reasons included increased hand washing/hand sanitizers, prolonged exposure to air-conditioning leading to excessive skin dryness and increased stress from being confined indoors and excessive drooling.

Supplementary Figure 1. Ten-point scale rating of adherence to treatment plan before and during lockdown period

Figure legend.

The figures show self-reported treatment adherence before (blue bars) and during (red bars) the lockdown period for subjects with Eczema (2A), Asthma (2B) and Allergic Rhinitis (2C). Subjects reported treatment adherence according to a ten-point rating scale where a score of 1 denoted poor adherence and 10 denoted perfect adherence.

Supplementary Table 1. Demographics of the study population.

Demographic Variable ⁺	Demographic Variable ⁺	All (n = 173)	Eczema (n = 115)	Asthma (n = 29)	Allergic rhinitis (n = 68)
Age (years), mean (SD)		7.1 (4.2)	6.3 (4.2)	7.8 (4.0)	8.6 (3.7)
Age groups (years)	0 – 3	30 (17.3)	28 (24.3)	2 (6.9)	2 (2.9)
	4 – 6	62 (35.8)	43 (37.4)	12 (41.4)	20 (29.4)
	7 – 12	60 (34.7)	34 (29.6)	11 (37.9)	35 (51.5)
	13 – 18	21 (12.1)	10 (8.7)	4 (13.8)	11 (16.2)
Gender	Male	89 (51.4)	55 (47.8)	15 (51.7)	41 (60.3)
	Female	84 (48.6)	60 (52.2)	14 (48.3)	27 (39.7)
Number of Allergic Diseases per subject	1	141 (81.5)			
	2	25 (14.5)			
	3	7 (4.0)			
Concomitant Allergic Disease	Eczema	115 (66.5)	-	11 (37.9)	22 (32.4)
	Asthma	29 (16.8)	11 (9.6)	-	13 (19.1)
	Allergic rhinitis	68 (39.3)	22 (19.1)	13 (44.8)	-
Caregiving arrangement during lockdown	Childcare facility	4 (2.3)	8 (7.0)	5 (17.2)	7 (10.3)
	Home-based	169 (97.7)	107 (93.0)	24 (82.8)	61 (89.7)
	Parent	130 (75.1)	88 (76.5)	17 (58.6)	48 (70.6)
	Others [^]	39 (22.5)	19 (16.5)	7 (24.1)	13 (19.1)

Demographic Variable ⁺	Demographic Variable ⁺	All (n = 173)	Eczema (n = 115)	Asthma (n = 29)	Allergic rhinitis (n = 68)
Primary caregiver for treatment administration*	Self-administered	9 (5.2)	6 (5.2)	3 (10.3)	3 (4.4)
	Parent	142 (82.1)	93 (80.9)	23 (79.3)	59 (86.8)
	Grandparent	15 (8.7)	13 (11.3)	1 (3.4)	3 (4.4)
	Domestic helper	17 (9.8)	9 (7.8)	2 (6.9)	9 (13.2)
	Not on regular medications	3 (1.7)	2 (1.7)	1 (3.4)	7 (10.3)

⁺ Reported as frequency (percentage) unless otherwise stated.

babysitter, patient

* Some subjects had more than one caregiver administering treatment

Supplementary Table 2. Patient-rated scores for allergic diseases

Score		Before lockdown	During lockdown	p-value
POEM (Max 28)	Overall score, median (IQR)	9 (3-13)	6 (3-10)	0.001
Individual questions	Itch	2 (1-4)	2 (1-3)	0.005
	Disturbed sleep	1 (0-2)	1 (0-2)	0.058
	Bleeding	1 (0-2)	0 (0-1)	0.002
	Weeping/oozing	0 (0-1)	0 (0-1)	0.079
	Cracked skin	1 (0-2)	0 (0-1)	0.017
	Flaking skin	1 (0-2)	0 (0-1)	0.003
	Rough and dry skin	2 (1-3)	2 (1-3)	0.394
ACT (Max 25)	Overall score, median (IQR)	22 (19-25)	23 (20-25)	0.063
Individual questions	Activity limitation	5 (4-5)	5 (4-5)	0.157
	Shortness of breath	5 (4-5)	5 (4-5)	1.00
	Nocturnal symptoms	5 (4-5)	5 (5-5)	0.143
	Use of rescue treatment	5 (4-5)	5 (4-6)	0.058
	Asthma control rating	4 (3-5)	5 (3-5)	0.097
TNSS (Max 12)	Overall score, median (IQR)	3 (2-4)	3 (2-5)	0.299
Individual questions	Nasal obstruction	1 (1-2)	1 (1-2)	0.655

Score		Before lockdown	During lockdown	p-value
Nasal itch/sneezing		1 (1-2)	1 (1-2)	0.097
Rhinorrhea		0 (0-1)	0 (0-1)	0.175

POEM: Patient Oriented Eczema Measure

ACT: Asthma Control Test

TNSS: Total Nasal Symptom Score

Bold p values indicate statistical significance ($p < 0.05$)



