

Children's Mercy Kansas City

SHARE @ Children's Mercy

Manuscripts, Articles, Book Chapters and Other Papers

6-2021

Digital Health Technology in Asthma: A Comprehensive Scoping Review.

Giselle Mosnaim

Guilherme Safioti

Randall Brown

Michael DePietro

Stanley J. Szefler

See next page for additional authors

Follow this and additional works at: <https://scholarlyexchange.childrensmercy.org/papers>



Part of the [Allergy and Immunology Commons](#)

Recommended Citation

Mosnaim G, Safioti G, Brown R, et al. Digital Health Technology in Asthma: A Comprehensive Scoping Review. *J Allergy Clin Immunol Pract.* 2021;9(6):2377-2398. doi:10.1016/j.jaip.2021.02.028

This Article is brought to you for free and open access by SHARE @ Children's Mercy. It has been accepted for inclusion in Manuscripts, Articles, Book Chapters and Other Papers by an authorized administrator of SHARE @ Children's Mercy. For more information, please contact library@cmh.edu.

Creator(s)

Giselle Mosnaim, Guilherme Safioti, Randall Brown, Michael DePietro, Stanley J. Szeffler, David M. Lang, Jay M. Portnoy, Don A. Bukstein, Leonard B. Bacharier, and Rajan K. Merchant

Digital Health Technology in Asthma: A Comprehensive Scoping Review



Giselle Mosnaim, MD, MS^a, Guilherme Safioti, MD^b, Randall Brown, MD, MPH^c, Michael DePietro, MD^c, Stanley J. Szeffler, MD^d, David M. Lang, MD^e, Jay M. Portnoy, MD^f, Don A. Bukstein, MD^g, Leonard B. Bacharier, MD^h, and Rajan K. Merchant, MD, FAAAAIⁱ Evanston, Ill; West Chester, Pa; Aurora and Cleveland, Colo; Kansas City, Mo; Greenfield, Wis; Nashville, Tenn; Woodland, Calif; and Amsterdam, The Netherlands

What is already known about this topic? Digital technology provides an opportunity to improve and individualize asthma self-management significantly across a variety of intervention types; however, the impact of different digital intervention characteristics has yet to be assessed.

What does this article add to our knowledge? Significant heterogeneity exists in study designs, patient populations, and outcomes measurement for digital interventions; more alignment is needed to measure impacts accurately on different dimensions of care and to guide future successful interventions.

How does this study impact current management guidelines? This scoping review does not directly affect current guidelines for asthma self-management, but it is hoped that it will inform the design of future digital intervention studies.

BACKGROUND: A variety of digital intervention approaches have been investigated for asthma therapy during the past decade, with different levels of interactivity and personalization and a range of impacts on different outcome measurements. **OBJECTIVE:** To assess the effectiveness of digital interventions in asthma with regard to acceptability and outcomes and evaluate the potential of digital initiatives for monitoring or treating patients with asthma. **METHODS:** We evaluated digital interventions using a scoping review methodology through a literature search and review. Of

871 articles identified, 121 were evaluated to explore intervention characteristics, the perception and acceptability of digital interventions to patients and physicians, and effects on asthma outcomes. Interventions were categorized by their level of interactivity with the patient. **RESULTS:** Interventions featuring non-individualized content sent to patients appeared capable of promoting improved adherence to inhaled corticosteroids, but with no identified improvement in asthma burden; and data-gathering interventions appeared to have little effect on adherence or asthma

^aDivision of Pulmonary, Allergy and Critical Care, Department of Medicine, North Shore University Health System, Evanston, Ill

^bTeva Pharmaceutical Industries, Ltd, Amsterdam, The Netherlands

^cTeva Branded Pharmaceutical Products R&D, Inc, West Chester, Pa

^dThe Breathing Institute and Pulmonary Medicine Section, Children's Hospital Colorado and University of Colorado School of Medicine, Anschutz Medical Campus, Aurora, Colo

^eDepartment of Allergy and Clinical Immunology, Respiratory Institute, Cleveland Clinic, Cleveland, Ohio

^fPediatric Allergy and Immunology, Children's Mercy Hospital, Kansas City School of Medicine, Kansas City, Mo

^gAllergy, Asthma and Sinus Center, Milwaukee, Greenfield, Wis

^hDepartment of Pediatrics, Monroe Carell Jr Children's Hospital at Vanderbilt University Medical Center, Nashville, Tenn

ⁱWoodland Clinic Medical Group, Allergy Department, Dignity Health, Woodland, Calif

Conflicts of interest: G. Mosnaim currently receives research grant support from Teva, Sanofi-Regeneron, Genentech, and Alk-Abello; received research grant support from AstraZeneca, GlaxoSmithKline, and Propeller Health; owned stock in Electrocore; and served as a consultant and/or member of a scientific advisory board for GlaxoSmithKline, Sanofi-Regeneron, Teva, Novartis, AstraZeneca, Boehringer Ingelheim, and Propeller Health. G. Safioti, R. Brown, and M. DePietro are employees of Teva Pharmaceuticals. S. J. Szeffler has consulted for AstraZeneca, Boehringer-Ingelheim, GlaxoSmithKline, Novartis, Propeller Health, Regeneron, and Sanofi; has received research support from the National Institutes of Health, the National Heart, Lung, and Blood Institute, Propeller Health, the Colorado Department of Public Health and Environment Colorado

Cancer, and Cardiovascular and Pulmonary Disease Program. D. M. Lang is currently past president of AAAAI; serves as a member of the Editorial Board for *Allergy and Asthma Proceedings*; is Topic Editor for DynaMed; is associate editor for *J Asthma*; and is delegate to National Quality Forum representing the American Academy of Allergy, Asthma & Immunology. J. M. Portnoy has received honoraria as a speaker from Thermo Fisher and has consulted for BioCryst. D. A. Bukstein has received honoraria as a speaker from AstraZeneca, Genentech, Novartis, Teva, Regeneron, and ALK. L. B. Bacharier reports personal fees from GlaxoSmithKline, Genentech, Novartis, Merck, DBV Technologies, Teva, Boehringer Ingelheim, AstraZeneca, WebMD/Medscape, Sanofi, Regeneron, Vectura, and Circassia; and receives royalties from Elsevier. R. K. Merchant has received honoraria from Teva, Sanofi, and AstraZeneca for participation in advisory boards; and has served as speaker and consultant for Propeller Health; and his spouse is an employee of and holds stock options in Horizon Therapeutics. Received for publication October 7, 2020; revised manuscript received and accepted for publication February 4, 2021.

Available online February 27, 2021.

Corresponding author: Michael DePietro, MD, Teva Branded Pharmaceutical Products R&D, Inc, 145 Brandywine Pkwy, West Chester 19380, Pa. E-mail: Michael.Depietro01@tevapharm.com.

2213-2198

© 2021 The Authors. Published by Elsevier Inc. on behalf of the American Academy of Allergy, Asthma & Immunology. This is an open access article under the CC BY license (<http://creativecommons.org/licenses/by/4.0/>).

<https://doi.org/10.1016/j.jaip.2021.02.028>

Abbreviations used

ACT- Asthma Control Test
 HCP- Health care professional
 ICS- Inhaled corticosteroids
 SABA- Short-acting β -agonist
 SMS- Short message service

burden. Evidence of improvement in both adherence and patients' impairment due to asthma were seen only with interactive interventions involving two-way responsive patient communication. Digital interventions were generally positively perceived by patients and physicians. Implementation was considered feasible, with certain preferences for design and features important to drive use.

CONCLUSIONS: Digital health interventions show substantial promise for asthma disease monitoring and personalization of treatment. To be successful, future interventions will need to include both inhaler device and software elements, combining accurate measurement of clinical parameters with careful consideration of ease of use, personalization, and patient engagement aspects. © 2021 The Authors. Published by Elsevier Inc. on behalf of the American Academy of Allergy, Asthma & Immunology. This is an open access article under the CC BY license (<http://creativecommons.org/licenses/by/4.0/>). (J Allergy Clin Immunol Pract 2021;9:2377-98)

Key words: Asthma; eHealth; Connectivity; Inhalers; Adherence; Monitoring; Disease control; Perception; Acceptability; Feasibility

INTRODUCTION

Rationale

Asthma continues to be a significant chronic disease burden, with estimated patient numbers approaching 350 million globally.¹ Asthma is a major source of direct and indirect economic cost in addition to reduced quality of life and premature death in patients of all ages, including children.² Several areas of unmet clinical need are associated with conventional asthma management; while medications such as inhaled short-acting β -agonists (SABA) and inhaled corticosteroids (ICS) are efficacious, considerable long-term commitment to asthma self-management on behalf of the patient is required for therapeutic success.

Asthma self-management includes advice and education on self-monitoring and a defined asthma management plan supported by regular professional review. Self-management is recognized as an effective method to improve asthma control and quality of life and reduce unscheduled consultations and hospitalizations across diverse demographic groups.³ For the patient, adherence to medication regimens is a key element of successful asthma management, in conjunction with wide-ranging factors including disease perception, patient education and understanding of symptoms, comorbidities, inhaler technique, access to medications, and health care professional (HCP) support.⁴ On the part of clinicians, implementation of evidence-supported management guidelines for assessment, monitoring, patient education, control of environmental factors, and pharmacologic treatment is a key factor, although adherence to such guidelines varies among different physician groups.^{5,6}

Several areas of asthma self-management could be improved using digital technology, including disease disparity, medication adherence, patient—clinician communication, personalized patient education, and passive monitoring of patient characteristics and behaviors that could prompt timely intervention. Technological innovations have the potential to support patients with disease self-management, allow for remote management by health care providers, and make future asthma management more proactive.⁷

Researchers have explored a variety of different digital asthma approaches including passive education, interactive websites, and electronic medication monitoring devices. Given the increasing prevalence of asthma worldwide and patient dependence on traditional inhaler devices to manage the disease, the inclusion of more advanced technology as a part of asthma self-management will be an essential next step to improving therapy for respiratory diseases. Lockdown and social distancing as a result of the COVID-19 pandemic have caused increased uptake and reliance on videoconferencing and telemedicine in various therapy areas.⁸ Increased familiarity with telemedicine provides an opportunity for remote monitoring and disease management, particularly in areas where access to HCPs is challenging.

Given the wealth of available literature and heterogeneity of digital technologies used to varying degrees of success, we performed a scoping review to assess the available research in this field. Scoping reviews provide a preliminary assessment of the size and scope of available literature on a topic and characterize the nature of published studies to provide an overview of large, heterogeneous bodies of literature. Furthermore, scoping reviews provide a platform for more precise systematic reviews to look at specific research questions. This scoping review therefore seeks to evaluate the different options that have been explored and assess the future utility of digital health technology in asthma.

Objective

This study aimed to evaluate the current and potential future usability, acceptability, uptake, effectiveness, and adoption of digital health technology for treating or monitoring asthma using a scoping review methodology. The potential future reach and durability of these technologies will also be considered.

METHODS

Scoping review approach

We conducted a systematic scoping review based on the methodology described⁹ and the Preferred Reporting Items for Systematic Reviews and Meta-Analyses extension for Scoping Reviews guidelines,¹⁰ including identifying the research question, identifying relevant studies, selecting the study, charting data, collating, and summarizing and reporting the results. The purpose of a scoping review is to give a broad overview of the available literature without being directed toward a single discrete research question.

Sources and searches

We searched PubMed, Embase, and the Cochrane Library for January 2010 to June 2019 using the search terms and criteria in [Table E1](#) (in this article's Online Repository at www.jaci-inpractice.org), adapted for each database, to identify asthma studies that assessed the effect of digital health technology interventions on patient outcomes, including symptoms, adherence, and medication use or therapy regimen. We also included studies that incorporated the acceptability of digital interventions to patients and HCPs and the

TABLE I. Classification of different intervention types

| Intervention category | Intervention types |
|-----------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| Generalized studies (not patient-specific or bidirectional): content given to patient | <ul style="list-style-type: none"> Automated/speech recognition and SMS dose and pharmacy-refill reminders Noninteractive education or motivational coaching by telephone, text, e-mail and other digital means (eg, simple access to static content) |
| Patient-specific, noninteractive studies (unidirectional): data collected from or sent to patient | <ul style="list-style-type: none"> Decision support tool for randomized controlled trial enrollment Device monitoring: noninhaler digital parameter tracking (eg, lung function) not resulting in immediate or ongoing asthma management feedback to patient (ie, data collection to inform health care professional at next visit) Individualized static education and/or motivation Digital questionnaires (by SMS/Web/phone) including ecological momentary association for data collection only (ie, not informing therapy) Inhaler only: data collected on device from digital inhalers or adherence monitors but not fed back to patient for self-management Inhaler platform: data collected from digital inhalers and a Web or app platform but not fed back to patient for self-management Platform only: online digital chat platform (by Web or mobile app) providing nonindividualized education |
| Patient-specific interactive studies including telehealth and video counseling: bidirectional interaction | <ul style="list-style-type: none"> Interactive education or motivational coaching Device monitoring: noninhaler digital parameter tracking (eg, lung function, inhaler technique) with feedback to patient (usually by smartphone app) Digital inhalers with electronic adherence monitors and individualized patient feedback Digital inhaler studies with accompanying individualized asthma management Web or app platform Interactive asthma management platforms that support self-management by collecting symptoms or Asthma Control Test/other data and give responses to guide patient's asthma treatment Telemedicine interventions by phone, video, or SMS |

SMS, short message service.

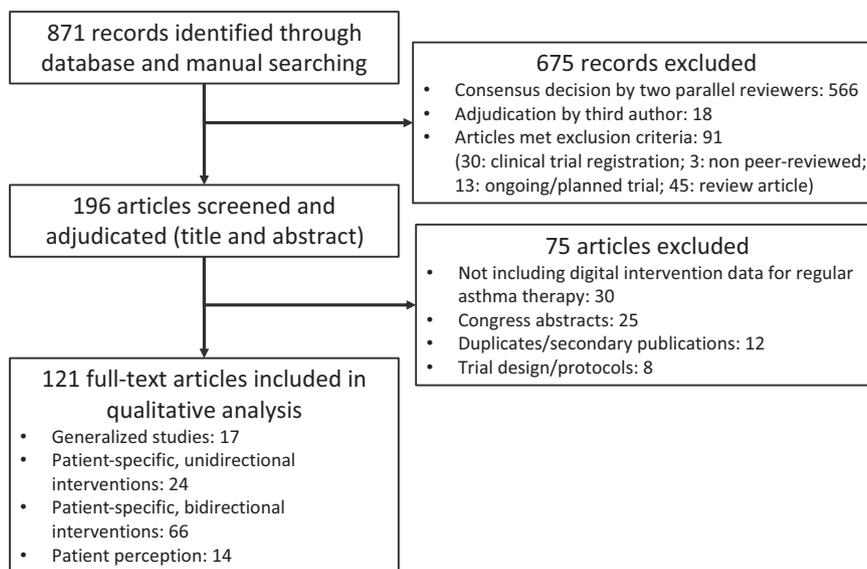


FIGURE 1. Articles selected for analysis.

feasibility of their wider uptake. Congress abstracts were excluded from the results. Database searches were supplemented by manual searches and references as appropriate.

Nonrelevant articles and duplicates were excluded; the titles and abstracts of remaining citations were reviewed by two authors (G.S. and G.M.) in parallel; a third author (R.K.M.) adjudicated discrepancies. Full-text articles were then obtained and a second round of screening was conducted (two reviewers worked in parallel and adjudication was performed by a third reviewer) and further articles were excluded as necessary. Articles were assessed for the type of

digital health technology employed, study design, size of the study, and outcomes assessed.

Data extraction and synthesis

The full text of the selected articles was used to identify the types of technology used. Interventions were grouped (Table I) according to their primary characteristic (in which interventions contained more than one aspect), specifically, according to their level of interactivity with the patient. Interventions in which non-individualized information was sent to patients on an automated

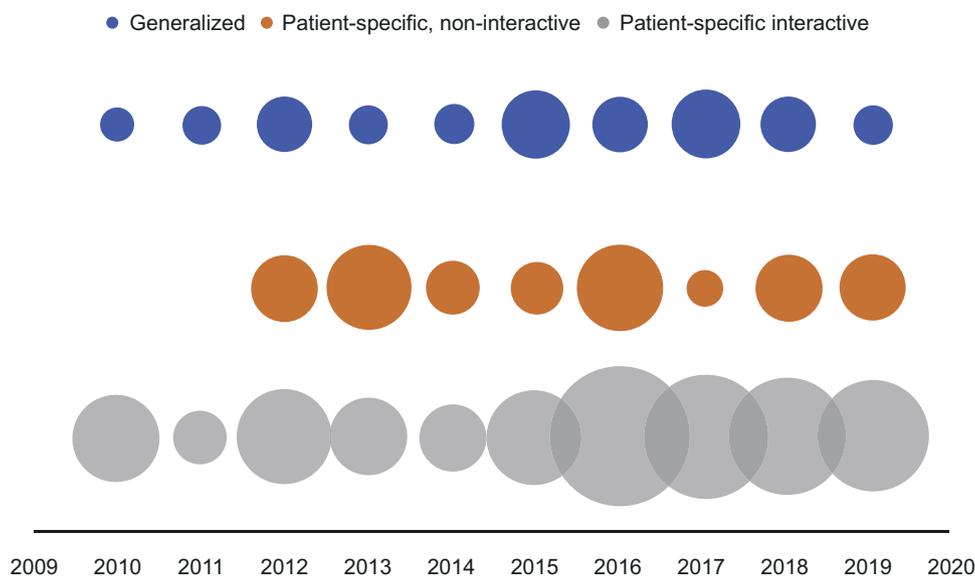


FIGURE 2. Bubble plot showing relative numbers of publications for different intervention categories by year.

basis (eg, short message service (SMS)-based dose reminders and automated pharmacy prescription refill calls, or access to noninteractive, non—patient specific digital educational material) were termed ‘generalized studies’. Interventions in which individualized information was gathered from or sent to patients (eg, parameter monitoring by a device with no feedback to the patient, or provision of patient-specific education) were termed ‘patient-specific, noninteractive studies’. Interventions composed of an interactive relationship with the patient, in which insights based on data collected from the patient or resulting from patient behavior were fed back to the patient to assist with asthma management on a personalized basis (eg, an asthma management platform responding to adherence data from a digital inhaler or questionnaire feedback from the patient) were termed ‘patient-specific interactive studies’. Within these broad intervention categories, studies were further grouped according to the type of technology involved (Table I).

Details of the studies performed (eg, subject characteristics, sample size, randomization) and outcomes, including adherence, asthma control or impairment and health care use were captured in a standardized table format as part of the data charting process. Adherence outcomes were assessed for being subjective (ie, reliant on questionnaire data or patient recollection) or objective (ie, monitored automatically as a result of medication use, including one study that calculated adherence from electronic medical records).

To assess the impact of the different intervention categories, we identified studies in which a positive effect (vs no effect or a negative effect) was reported with the intervention in three domains: adherence to ICS (and long-acting β_2 -agonist) therapy, impairment caused by asthma (most commonly assessed by Asthma Control Test [ACT] score, but including changes in symptoms and quality of life when reported), and health care resource use (urgent consultations, emergency department visits, or hospitalizations). Because of heterogeneity in data reporting, to include as many studies as possible in our broad overview, we considered statistically significant, numerical, and anecdotal reports of benefit as evidence of positive effects in

each of these three domains. Counts for studies reporting positive versus no effect or negative effects were then summarized.

Articles that reported patient and HCP perceptions of digital technologies were assessed separately, and relevant observations were extracted.

RESULTS

Selection of sources of evidence

Our selection process identified 121 articles for inclusion in this scoping review (Figure 1); 17 articles described generalized studies, 23 described patient-specific, noninteractive interventions, and 66 described patient-specific interactive interventions. Figure 2 shows a bubble plot of the number of articles for each intervention category published each year. During the past 5 years, interactive interventions were investigated more frequently than noninteractive initiatives, possibly because of the increased availability of mobile Internet access and personal smart devices. Fourteen articles were identified that investigated patients’ and clinicians’ perceptions of digital interventions.

Table II lists the number of studies reporting positive effects on outcomes for different intervention types.

Generalized studies

There were 17 generalized interventions, eight of which employed reminders or automated phone calls for medication doses or pharmacy fills, and nine that provided asthma education or asthma management motivation ($n = 9$). Most were randomized ($n = 12$) and most had an intervention period of 3 or more months ($n = 15$) (Table III).

Six of the studies described a change in adherence with the intervention; all of these reported a positive effect (6 positive vs 0 no effect or negative). Adherence in these studies was measured using both subjective ($n = 3$) and objective ($n = 3$) approaches.

TABLE II. Outcomes reporting for intervention categories and types

| Intervention | Articles, n | Adherence | | Asthma impairment | | Health care use | |
|--------------------------------------------------|-------------|-----------|-----------|-------------------|-----------|-----------------|-----------|
| | | Positive | No effect | Positive | No effect | Positive | No effect |
| Generalized studies | 17 | 6 | 0 | 3 | 7 | 0 | 5 |
| Reminders and pharmacy fill | 8 | 6 | 0 | 0 | 5 | 0 | 3 |
| EM | 9 | 0 | 0 | 3 | 2 | 0 | 2 |
| Patient-specific, noninteractive, unidirectional | 24 | 2 | 4 | 2 | 4 | 0 | 2 |
| Decision support | 1 | 0 | 0 | 1 | 0 | 0 | 0 |
| Digital questionnaire | 9 | 0 | 1 | 1 | 1 | 0 | 1 |
| Device monitoring | 8 | 0 | 1 | 0 | 1 | 0 | 0 |
| EM | 2 | 1 | 1 | 0 | 0 | 0 | 0 |
| Inhaler only | 2 | 1 | 0 | 0 | 1 | 0 | 0 |
| Inhaler platform | 1 | 0 | 0 | 0 | 0 | 0 | 0 |
| Platform only | 1 | 0 | 1 | 0 | 1 | 0 | 1 |
| Patient-specific interactive, bidirectional | 66 | 17 | 8 | 31 | 15 | 12 | 14 |
| EM | 2 | 0 | 1 | 1 | 1 | 0 | 1 |
| Inhaler only* | 2 | 1 | 0 | 1 | 0 | 0 | 1 |
| Inhaler platform | 14 | 5 | 0 | 5 | 2 | 3 | 0 |
| Platform only | 32 | 9 | 6 | 15 | 9 | 5 | 8 |
| Telemedicine | 14 | 2 | 1 | 8 | 3 | 4 | 3 |
| Device monitoring | 2 | 0 | 0 | 1 | 0 | 0 | 1 |

EM, education and/or motivation.

*The study by Chan et al¹¹ was not included in positive/negative calculations because it was the same study as Chan et al.¹²

Few studies of generalized interventions reported a positive effect on asthma impairment (3 positive vs 7 no effect or negative), and none reported meaningful improvements in health care use (0 positive vs 5 no effect or negative) (Table II and Figure 3).

Patient-specific, noninteractive, unidirectional interventions

Of the 24 studies using patient-specific, noninteractive, unidirectional interventions, most focused on either digital questionnaire-based studies (n = 9) or device monitoring studies (without feedback to the patient; n = 8). Thirteen studies were randomized and 11 were single-arm. Nine interventions lasted for less than 3 months (Table IV).

Overall, few studies reported positive effects; only two studies found benefit in each of the adherence and asthma impairment domains and none reporting improvements in health care use (Table II and Figure 3). Positive effects on adherence were noted in one study of education provision and one involving adherence tracking with an automated reminder call; moreover, positive effects on asthma impairment resulted from one study involving a smartphone symptom and ACT diary with additional education and motivation, and one study in which a decision support algorithm was used to enter patients into different randomized controlled trials.

Patient-specific interactive, bidirectional interventions

Patient-specific, bidirectional interactive interventions were the largest group of interventions (n = 66). Most often, these entailed an asthma management platform (n = 32), a digital inhaler combined with an asthma management platform (n = 14), or a telemedicine-based intervention (n = 14). Most studies (n = 45) were randomized, and 15 of the interventions

lasted less than 3 months (including two with the study period not stated) (Table V).

Reported positive effects outweighed no effect for both adherence (17 vs 8 studies, respectively) and asthma impairment (31 vs 15 studies, respectively) (Table II and Figure 3). Adherence was measured subjectively (n = 18) and objectively (n = 14) in a similar number of studies. Twelve studies found reductions in health care use, although a similar number (n = 14) found no effect.

This was the only intervention category within which improvements in both adherence and asthma impairment were reported more often than not and were seen consistently across the three most-studied intervention types.

Acceptability and feasibility of interventions

When reported, all types of digital interventions were considered to be acceptable or favorable to patients or their caregivers and feasible to implement, although this aspect was not rigorously investigated in articles reporting intervention results.^{13,14,16-19,22-24,26,28,43,56,87,105,108,109,111,115,117} Digital inhaler systems were described as feasible to implement^{31,61,62,65,69-71} and acceptable to most patients.^{11,31,61,70,118} Similarly, digital asthma management platforms were reported to be both feasible^{52,57,82,84,93,119} and acceptable.^{75,76,92,96,100-102,107} One study reported low use of a Web portal that was not specifically targeted to symptomatic asthma patients.⁸¹

Although patients were generally satisfied with digital devices and were willing to use them, alerts were sometimes perceived as unwelcome if they were received at inopportune times.³⁵ Non-inhaler devices (including Fitbits (Google LLC, San Francisco, CA) to track activity levels and sleep, and devices designed to assess lung function) were also considered acceptable by patients.^{35,52,54}

TABLE III. Generalized studies

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|------------------------|------|------------------------|-------------------|-------------------------------------------------------|------------|--------------|-----------------------------------------------------------------------------|------------------------------------------------------------|------------------------------------------------------------------------------|
| 13 | Adults only | 4510 | NS | Reminder | Refill | SA | 2 y | Significant but small improvement | — | No change in ED visits or hospitalizations |
| 14 | Adolescent or younger | 1187 | NS | Reminder | Refill | R | 24 mo | Significant improvement in ICS adherence at 24 mo | — | No difference in ED visits, primary care visits or hospitalizations |
| 15 | Adults only | 100 | NS | Reminder | SMS reminder | SA | 5 y | — | No difference in ACT scores between groups | — |
| 16 | Adolescents and adults | 64 | NS | Reminder | SMS reminder | R | 6 mo | 2.75% increase in adherence each month, but improvements were not sustained | No significant difference between arms | ED visits and hospitalizations for asthma were rare |
| 17 | Adolescent or younger | 89 | NS | Reminder | SMS reminder plus static education | R | 3 wk | Significant improvement in self-reported 7-d adherence | No significant change in ACT | — |
| 18 | Adults only | 26 | Mild, moderate, severe | Reminder | SMS reminder | R | 12 wk | Significant increase vs control | — | — |
| 19 | Adolescents and adults | 12 | NS | Reminder | SMS reminder | R | 3 mo | — | No significant change in mean self-reported asthma control | — |
| 20 | Adults only | 8517 | NS | Reminder | Refill | R | 18 mo | Modest but significant increase in ICS adherence | No significant change in asthma control | No significant effect on acute asthma health care use |
| 21 | Adolescent or younger | 359 | NS | EM | Static education to prompt physician questioning | R | 1 y | — | — | — |
| 22 | Adults only | 655 | NS | EM | Static education by DVD for breathing retraining | R | 12 mo | — | No significant improvement in symptom scores | — |
| 23 | Adults only | 150 | NS | EM | SMS education | R | 13 mo | — | — | No significant difference in incidence of ED visits between groups |
| 24 | Adolescent or younger | 64 | NS | EM | Static EM, updated at intervals | R | 3 mo | — | Significant increase in cACT | — |
| 25 | Adults only | 330 | NS | EM | Static education to encourage written asthma plan use | R | 12 mo | — | No statistically significant change in ACQ scores | No significant change in ED or non-emergency health care professional visits |

| | | | | | | | | | | |
|----|-----------------------|-----|----|----|---------------------------------------------------|----|----------|---|-----------------------------------------------------------------------|---|
| 26 | Adults only | 144 | NS | EM | Static video education during face-to-face visits | SA | 1 mo | — | Significant improvement in ACT score | — |
| 27 | Adults only | 34 | NS | EM | Static Web-based education | SA | 1 y | — | — | — |
| 28 | Adolescent or younger | 50 | NS | EM | Static Web-based education | SA | 2 wk | — | — | — |
| 29 | Adults only | 51 | NS | EM | Static Web-based education | R | 3 months | — | Significant improvements in self-reported asthma symptoms and control | — |

(c)ACT, (childhood) Asthma Control Test; ACQ, Asthma Control Questionnaire; ED, emergency department; EM, education and/or motivation; ICS, inhaled corticosteroid; SA, single-arm; NS, not specified; R, randomized; SMS, short message service.

Cost-effectiveness

Only one¹¹² of the five studies to comment on cost-effectiveness^{13,22,109,112,116} found that the digital intervention was not cost-effective, although formal investigations of cost-effectiveness were not performed. A smart nebulizer system was associated with decreased costs of additional treatments associated with asthma, such as antibiotics and corticosteroids.⁶⁴ Web-based monitoring of asthma control was deemed likely to be cost-effective after an analysis of an economic model.³⁹

Perceptions of digital health technology

Fourteen articles commented on perceptions regarding digital interventions,^{95,119-131} and all intervention types were widely considered to be acceptable and useful across groups of patients, prescribers, and families.

Patients' uptake of smartphone apps was affected by perceived need (ie, favorable if asthma not considered to be well-controlled)¹²⁵; for smartphone app uptake in general, ease of access (ie, downloading) was considered important. Although social media could attract many initial downloads, long-term engagement was more successful with face-to-face recruitment by practices.¹²⁴

Adolescents thought that smartphone apps should be designed to be visually simple, easy to use, informative, and customizable, and include familiar components (eg, alerts that appeared and sounded like their usual messaging apps), and they were open to using apps for support and information.^{121,122,130,131} They preferred time-based to event-based reminders⁴⁴ and favored video and peer-chat functionality in addition to medication reminders.¹¹⁹ They were also sensitive to design choices that were too childish for their age range, but appreciated gamification of asthma management.¹²¹⁻¹²³

Adolescent patients and pharmacists had a positive attitude toward interactive mobile health interventions,¹¹⁹ and adults and adolescents were receptive toward educational and directive SMS messaging; adolescents were also in favor of sending messages to a support person (ie, parent or guardian).¹²⁵ Tablet- and smartphone-based asthma questionnaires were acceptable to patients and useful in certain situations (eg, when HCP time was limited, or in rural areas).¹²⁶ Patients and caregivers were supportive of interactive Web-based education provision based on patient information and preferences, and of augmented reality inhaler technique training.¹²⁸

Patients considered that smart devices and inhalers should be easy to use and convenient in terms of size and portability.^{95,127}

Clinicians valued the concept of intervisit improvement of adherence and lung function data and considered adherence data to be especially important, with a preference for ICS adherence data. Rescue inhaler information was considered valuable to access if patients were overusing them, because this could indicate a need to adjust the ICS regimen. The opportunity to use a patient's data to illustrate how adherence affected symptoms or impairment during consultations was also appreciated.¹²⁰

Some interesting concerns for clinicians included liability for acting on real-time data collection between visits when they could not respond, a preference to see data only before patient visits, and validation of data accuracy and security when devices could be shared.^{120,132}

DISCUSSION

In this scoping review, we surveyed a wide variety of approaches to implementing digital technology to improve patient

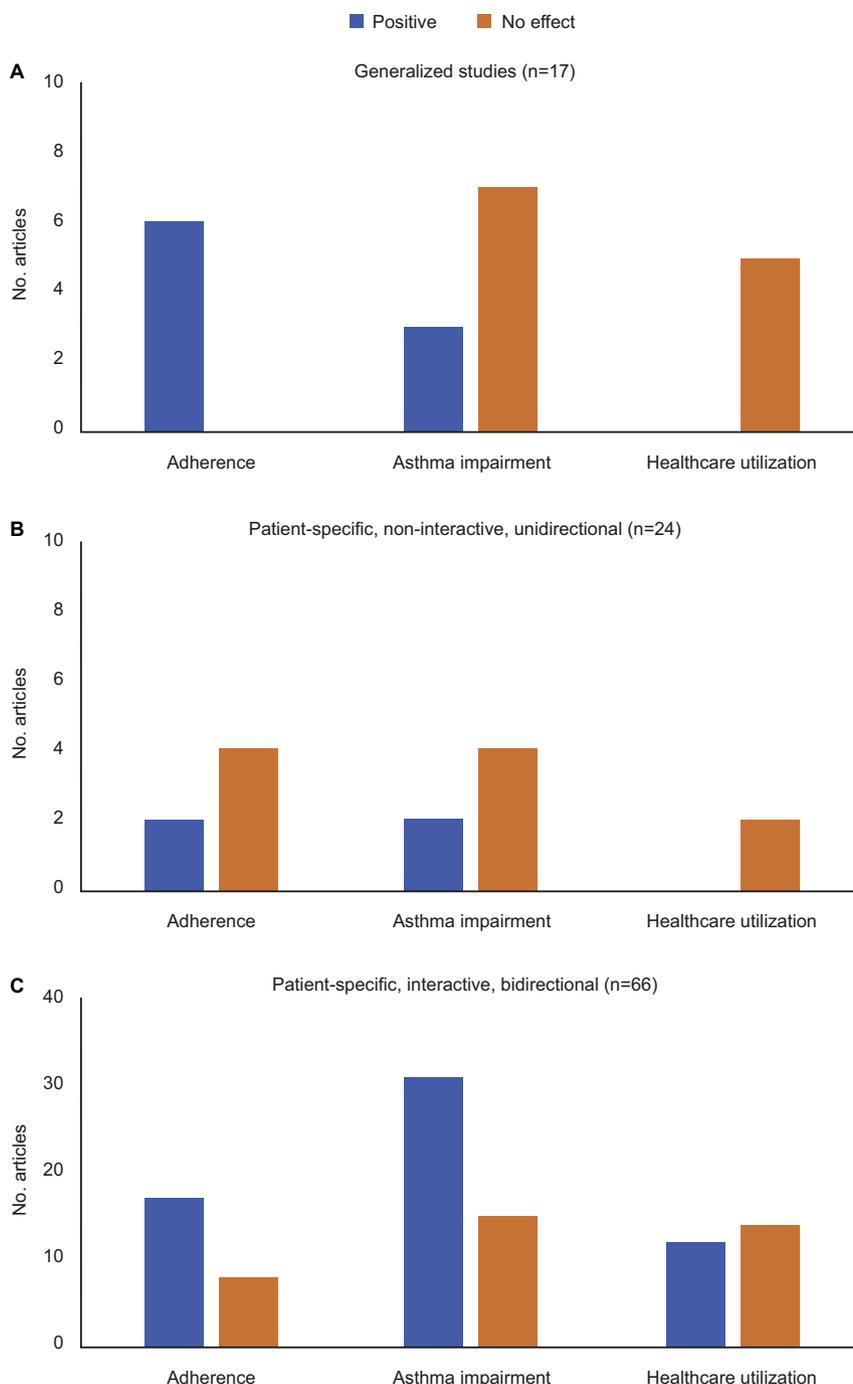


FIGURE 3. Balance of articles reporting positive effect versus no effect on adherence, asthma impairment, and health care use: A Generalized studies (n = 17). B Patient-specific, noninteractive, unidirectional interventions (n = 24). C Patient-specific interactive, bidirectional interventions (n = 66).

outcomes from asthma therapy. Such interventions usually focus on improving patient’s adherence to their treatment regimen, through dose and prescription-fill reminder systems, behavior change as a result of education and motivation, or increased access to health care professionals and asthma advice through telemedicine or interactive asthma management platforms. We considered the interventions identified by our literature search based on their interactivity with patients, and whether data

communication was unidirectional (data sent either from or to the patient) or bidirectional (data received from the patient prompting a response from the intervention). In terms of study outcomes, we looked for evidence of improvement with the various interventions in terms of patients’ adherence to ICS regimens, improvements in impairment owing to asthma, and improvements in health care use (by reductions in unplanned or emergency consultations).

TABLE IV. Patient-specific, noninteractive, unidirectional interventions

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|-----------------------|------|-----------------|-----------------------|----------------------------------------------------------------------------------------------------------|------------|--------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------|--------------------------------------------------------------------|
| 30 | Adolescent or younger | 1143 | NS | Decision support | Computerized randomized controlled trial decision support | R | 16 wk | Change in adherence during therapy not reported | Significant increase in proportion of participants gaining good control | — |
| 31 | Adolescent or younger | 41 | NS | Device monitoring | Propeller Health platform only for data collection | R | 30 d | Similar average daily adherence between intervention and control groups | Mean change in cACT similar between intervention and control groups | — |
| 32 | Adults only | 239 | Severe | Device monitoring | INCA adherence and inhaler technique only for data collection | SA | 3 mo | Change in adherence during therapy not reported | — | — |
| 33 | Adults only | 184 | NS | Device monitoring | INCA adherence and inhaler technique for periodic assessment | R | 3 mo | Change in adherence during therapy not reported; adherence via dose counter was significantly lower than by objective monitoring | Adherence was predictive of asthma exacerbations | — |
| 34 | Adults only | 200 | NS | Device monitoring | Smart spirometer only for data collection | SA | 2 mo | — | — | — |
| 35 | Adolescent or younger | 32 | NS | Device monitoring | PEFR monitoring by SMS and digital peak flow meter, cold symptom monitoring by SMS | SA | 3 wk | — | — | — |
| 36 | Adults only | 111 | NS | Device monitoring | Adherence monitor only for data collection | R | 24 wk | Change in adherence during therapy not reported; consistent overestimation of self-reported adherence vs digitally monitored | — | — |
| 37 | Adolescent or younger | 90 | NS | Device monitoring | Adherence monitor only for data collection | SA | 3 mo | Change in adherence during therapy not reported, ethnicity correlated with adherence | — | — |
| 38 | Adults only | 123 | NS | Device monitoring | INCA adherence and inhaler technique only for data collection | SA | 2-4 wk | Overall change in adherence during therapy not reported; significant correlation between adherence and socioeconomic class, education. Inclusion of inhaler technique errors significantly reduced adherence by dose counter | — | — |
| 39 | Adolescent or younger | 272 | NS | Digital questionnaire | Cost-effectiveness data for Lv et al ²³ ; Web-based ACT used to adjust treatment periodically | R | 4 mo | — | ACT used to adjust treatment | — |

(continued)

TABLE IV. (Continued)

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|------------------------|------|------------------------|-----------------------|----------------------------------------------------------------------------------------------|------------|--------------|---------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------|
| 40 | Adolescent or younger | 272 | NS | Digital questionnaire | See also Beerthuis et al ³⁹ ; Web-based ACT used to adjust treatment periodically | R | 1 y | — | No significant change in ACT in Web intervention group | No significant differences between groups in hospitalizations or steroid courses |
| 41 | Adolescent or younger | 84 | NS | Digital questionnaire | ADAM iPod cough monitoring and smartphone asthma diary for data collection only | SA | 7 d | — | — | High cough counts were associated with higher use of health care services, hospitalizations, and office visits |
| 42 | Adolescent or younger | 53 | NS | Digital questionnaire | Automated telephone EMA | SA | 1 mo | Change in adherence during therapy not reported | Change in ACT during therapy not reported | — |
| 43 | Adolescents and adults | 410 | NS | Digital questionnaire | ACT by SMS | R | 5 ± 1 wk | — | Significant improvements in ACT were detectable using SMS reporting | — |
| 44 | Adults only | 16 | NS | Digital questionnaire | EMA by SMS with automated prompts for data collection | SA | 2 wk | Change in adherence during therapy not reported | Change in symptoms during therapy not reported | — |
| 45 | Adolescent or younger | 228 | NS | Digital questionnaire | Web-based symptom diary for data collection only | R | 4 wk | Change in adherence during therapy not reported | Change in ACT and cACT score during therapy not reported | — |
| 46 | Adolescent or younger | 50 | Moderate | Digital questionnaire | Smartphone app symptom diary plus EM | R | 3 mo | No significant difference in self-reported medication adherence between groups with additional education/motivation | Significant improvement in cACT | — |
| 47 | All ages | 7593 | Mild, moderate, severe | Digital questionnaire | Regular data surveys from asthma patients | SA | 6 mo | Change during study not reported | Change during study not reported | — |
| 48 | Adolescent or younger | 68 | NS | EM | Individualized static education | R | 13 wk | No significant difference in adherence at wk 5 and 10 | — | — |
| 49 | Adolescents and adults | 216 | NS | EM | Tailored static EM | R | 9 mo | Significant improvement in adherence but less than clinically relevant level | — | — |

| | | | | | | | | | | |
|----|-----------------------|----|----|------------------|-------------------------------------------------------------------------------------------|----|-----------|------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------|------------------------------------------|
| 50 | Adolescent or younger | 81 | NS | Inhaler only | Adherence monitor for data collection only | SA | 12 mo | Change in adherence during therapy not reported; median adherence was 87% and relatively constant in most patients | Asthma control during 12-mo study period was high and lung function was normal | — |
| 51 | Adults only | 50 | NS | Inhaler only | Automated interactive call plus electronic dose tracker | R | 10 wk | 32% improvement in adherence over 10 wk | No difference in ACT | — |
| 52 | Adolescent or younger | 82 | NS | Inhaler platform | kHealth platform and digital phenotyping; data unidirectional; data available only to HCP | SA | 1 or 3 mo | Effect of intervention on controller adherence during study not reported; poor adherence associated with poor asthma control | Effect of intervention on ACT during study not reported; poor adherence associated with poor asthma control | — |
| 53 | Adults only | 67 | NS | Platform only | WeChat platform with education and HCP interaction | R | 3 mo | No significant difference in medication adherence between groups | No significant difference in improvement of asthma control between intervention and control groups | No significant difference between groups |

(c)ACT, (childhood) Asthma Control Test; EM, education/motivation; EMA, ecological momentary association; HCP, health care professional; ICS, inhaled corticosteroid; PEFR, peak expiratory flow rate; SA, single-arm; NS, not specified; R, randomized; SMS, short message service.

Generalized dose and prescription reminders and non-personalized education and motivation were effective in improving adherence, but not impairment owing to asthma. Although subjective measurement of adherence is thought to overestimate adherence compared with objective measurement,^{33,36} both types of adherence measurement were employed across these generalized studies; it is possible that adherence improvements were not sustained long enough to make a meaningful impact on asthma impairment. The patient-specific, noninteractive interventions included several different intervention types, but few reported positive impacts on either adherence or asthma impairment. However, the most-studied category of intervention, patient-specific interactive bidirectional interventions, reported the greatest number of improvements in adherence and asthma impairment. These benefits were largely found in studies of the most interactive interventions, those using interactive telemedicine, or those using an interactive management platform, with or without a digital inhaler. Each of these intervention types provides a high level of interactivity and instant feedback for patients through either live HCP consultation (telemedicine) or responses to monitored data or questionnaires (digital platforms with or without digital inhalers). Correlating patients' own asthma experiences (through ACT scores or other measured parameters) with their asthma management behaviors (eg, ICS adherence, trigger avoidance) may be a powerful engagement tool to motivate and educate patients about the need to stay on top of managing their condition.

We were able to identify broad trends in adherence and changes in asthma impairment among different intervention categories. The included studies generally did not find differences in health care use with interventions, either because it was not studied or because the incidence of emergency department visits or other unscheduled consultations was generally low.

All types of digital health interventions were well-received and acceptable to patients, caregivers, and clinicians. It is intuitive to think that digital interventions would be easily scalable to large populations and therefore be cost-effective, but currently there are few formal cost-effectiveness studies to confirm this assumption. In particular, the translation of improved adherence to improved symptom control and reduced health care use is not guaranteed, although several studies noted reductions in reliever or oral steroid use and reductions in unscheduled primary care visits. There may also be difficulties in transitioning coding and billing practices to accommodate digital interventions.

The progression of digital health monitoring and associated interventions in asthma still faces significant challenges. It is clear that a successful intervention must be able to combine objective monitoring with active patient engagement and education or motivation to maintain behaviors helpful to asthma therapy, and we have yet to develop an ideal system in this respect. Although there is further opportunity to incorporate digital interventions into patient care with respect to linking to patients' electronic health records and using data monitoring to trigger further HCP interventions, there is also the possibility of providing regular patient support through education about inhaler technique, avoidance of asthma triggers, and environmental monitoring and advice (eg, warnings related to high pollen count or air pollution), which could be equally important in reducing asthma burden.

The extensive implementation of digital interventions is attractive from the perspective of real-time population health monitoring as well as the increased opportunity and ability to

TABLE V. Patient-specific interactive, bidirectional interventions

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|-----------------------|-----|-----------------|-------------------|---------------------------------------------------------------------------------------------|------------|--------------|---------------------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------------------------------------------------|----------------------------------------------------------------------------------|
| 54 | Adolescent or younger | 22 | NS | Device monitoring | Vortex whistle and smartphone app for PEV | SA | 1 mo | — | — | — |
| 55 | Adults only | 72 | All | Device monitoring | FEV measurement with smartphone asthma monitoring app | R | 6 mo | — | Significant improvement in asthma control | No significant differences in unscheduled health care visits |
| 56 | Adults only | 44 | NS | EM | Individualized education in response to assignments | SA | 6 wk | — | Significant increase in mean ACT score (greater than MID) at 3 mo after intervention | — |
| 57 | Adults only | 51 | Mod | EM | Interactive website for education and improving adherence | R | 3 mo | No significant difference between groups | No significant difference in ACQ between groups | No significant difference between groups in HCP contact or steroid prescriptions |
| 11 | Adolescent or younger | 220 | NS | Inhaler only | Reliability data for Chan et al ¹² ; Adherence monitor with or without reminders | R | Not stated | — | Significant perceived and objective improvement in asthma control in conjunction with Chen et al ⁹⁰ | — |
| 12 | Adolescent or younger | 220 | NS | Inhaler only | See also Chan et al ¹¹ ; adherence monitor with or without reminders | R | 6 mo | Significant improvement in adherence with the reminder function in poorly adherent population at baseline; adherence then decline over 6 mo | Significant improvement in cACT vs control | No difference in frequency of ED visits |
| 58 | Adults only | 39 | NS | Inhaler platform | Adherence monitor and AMP platform | R | 3 mo | Baseline adherence and adherence vs control not reported. Slight reduction in adherence in intervention group during study | Greater improvement from baseline to 3 mo for intervention compared with control group | — |
| 59 | All ages | 495 | NS | Inhaler platform | Propeller Health platform monitoring of SABA use | R | 12 mo | — | Significant improvements in ACT in initially uncontrolled adults only; significantly larger decrease in SABA use vs routine care | — |
| 60 | Adolescent or younger | 77 | NS | Inhaler platform | Adherence monitor with periodic feedback | R | 12 mo | Significantly higher adherence in intervention group over 12 mo | No significant difference in improvement of asthma control between intervention and control groups | Significantly fewer hospital admissions in intervention group |

| | | | | | | | | | | |
|----|------------------------|-----|--------------------|------------------|-------------------------------------------------------------------|----|------------|----------------------------------------------------------------------------------------|---------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------|
| 61 | Adolescent or younger | 14 | Moderate to severe | Inhaler platform | Adherence monitor with motivational interviewing and feedback | SA | 3 mo | Change in adherence during therapy not reported | Borderline significant improvement in ACT ($P = .05$) | — |
| 62 | Adolescent or younger | 7 | NS | Inhaler platform | Adherence monitor and smartphone app | SA | 12 wk | Not performed owing to low patient numbers | — | — |
| 63 | All ages | 224 | NS | Inhaler platform | Propeller Health platform monitoring of SABA and ICS use | SA | 1 y | — | — | Significant reduction in asthma-related ED visits, and combined ED and hospitalization events |
| 64 | Adolescent or younger | 65 | NS | Inhaler platform | Smart nebulizer with smartphone app for HCP contact | R | 12 wk | Statistically significantly higher adherence at all time points vs control | — | Significantly lower frequency of ED visits, respiratory infections, antibiotics or systemic steroid use, and therapeutic cost vs control |
| 65 | Adolescent or younger | 12 | NS | Inhaler platform | Adherence monitor with smartphone app | SA | Not stated | Clinically significant improvement during pilot study | — | — |
| 66 | Adults only | 30 | NS | Inhaler platform | Adherence monitor for SABA use and interactive Web interface | SA | 4 mo | — | Small but statistically significant improvement in ACT scores | — |
| 67 | Adolescents and adults | 120 | NS | Inhaler platform | Adherence monitor for SABA use and interactive Web interface | SA | 1 mo | Rescue medication monitored rather than ICS | Significant reduction in SABA use during study | — |
| 68 | Adolescent or younger | 12 | NS | Inhaler platform | Adherence monitor and smartphone app | SA | 8 wk | Patients meeting target ICS adherence ($\geq 50\%$) increased from 8% to 58% at wk 8 | 58% of patients reached MID for ACT scores | — |
| 69 | Adolescents and adults | 143 | Moderate to severe | Inhaler platform | Adherence monitor and Web interface | R | 6 mo | Significant increase in adherence in groups receiving reminders and feedback | No significant difference in ACT between groups | — |
| 70 | Adults only | 32 | NS | Inhaler platform | Adherence monitor for data collection with customizable reminders | SA | 3 wk | Change in adherence during therapy not reported | — | — |
| 71 | All ages | 497 | NS | Inhaler platform | Adherence monitor with interactive smartphone app | SA | 12 mo | Rescue medication monitored rather than ICS | Significant increase in symptom-free days over 12 mo | — |

(continued)

TABLE V. (Continued)

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|-----------------------|-----|------------------|-------------------|-----------------------------------------------------|------------|--------------|---------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------|
| 72 | Adults only | 138 | NS | Platform only | breathe mobile health platform | R | 12 mo | Self-reported controller use increased before clinic visits | Post hoc analysis associated high ACT scores with app use | — |
| 73 | Adults only | 408 | NS | Platform only | Online asthma care tool | R | 12 mo | No significant difference between groups | Significant improvement in total ACT score with intervention | No significant difference between groups |
| 74 | Adults only | 60 | NS | Platform only | Interactive smartphone app | SA | 4 mo | — | Significant increase in ACT | — |
| 75 | Adults only | 100 | NS | Platform only | Online MAP asthma educational and monitoring portal | R | 6 mo | — | ACT evaluated but not reported separately | No significant difference in hospitalizations |
| 76 | Adults only | 327 | All | Platform only | POPET interactive smartphone app | R | 6 mo | — | Significant improvements in ACT and proportion of well-controlled patients | Significant reduction in unplanned hospital visits and ED visits |
| 77 | Adolescent or younger | 152 | Mild to moderate | Platform only | Asthma management smartphone app | R | 12 mo | Statistically significant higher treatment adherence vs control group | Statistically significantly higher cACT scores and fewer exacerbations vs control | — |
| 78 | Adolescent or younger | 193 | NS | Platform only | AsthmaCare smartphone asthma management app | R | 6 mo | — | — | Significant reduction in urgent care visits, but not ED visits or hospitalizations |
| 79 | Adolescent or younger | 34 | NS | Platform only | Smartphone AMP | R | 6 mo | — | Overall, no changes in ACT and self-efficacy scores between groups; adolescents with uncontrolled asthma had improved ACT within intervention group | — |
| 80 | Adults only | 103 | NS | Platform only | Online asthma patient community | R | 9 wk | No change in self-reported adherence | — | — |
| 81 | Adolescent or younger | 237 | All | Platform only | MyAsthma online asthma management portal | R | 12 mo | Significant increase in asthma medication changes or refills for children with uncontrolled asthma after parental use of portal | — | Significantly more medication changes and primary care asthma visits with portal use |
| 82 | Adults only | 44 | NS | Platform only | snuCare smartphone AMP and PFM | R | 8 wk | Significant improvement in adherence | — | — |

| | | | | | | | | | | |
|----|-----------------------|-----|--------------------|---------------|-----------------------------------------------------------------|----|------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------|
| 83 | Adults only | 48 | NS | Platform only | Interactive motivational interviewing and EMA | R | 3 mo | Trend for fewer doses missed vs control, but not statistically significant | Trend for improved ACT vs control, but not statistically significant | — |
| 84 | Adolescent or younger | 58 | NS | Platform only | Interactive smartphone app | R | 6 mo | Significant improvement in patients with low (<75%) baseline adherence | — | No significant change in ED or acute visits during study |
| 85 | Adolescent or younger | 20 | NS | Platform only | Smartphone AMP | SA | 8 wk | — | Significant increase in ACT for participants with uncontrolled asthma at baseline | — |
| 86 | Adults only | 21 | Moderate to severe | Platform only | Online P'ASMA asthma management portal | R | 4 wk | No significant difference between groups | No significant difference between groups | — |
| 87 | Adolescent or younger | 26 | Moderate to severe | Platform only | EM by phone plus tailored SMS | R | 3 mo | Reduced barriers to adherence | Clinically meaningful improvement in symptoms with medium to large effect sizes over 3 mo | — |
| 88 | Adults only | 89 | Moderate to severe | Platform only | Smartphone asthma diary with management feedback | R | 6 mo | — | Asthma symptom scores used to adjust therapy | Significantly fewer patients with unscheduled ED visits |
| 89 | Adults only | 120 | NS | Platform only | SMS monitoring with HCP feedback as necessary | R | 3 mo | — | No significant difference between groups | No significant difference between groups |
| 90 | Adults only | 424 | NS | Platform only | SMS monitoring with HCP feedback as necessary | R | 3 mo | — | No significant difference between groups | No significant difference in ED visits or hospitalizations between groups after 5 wk |
| 91 | Adults only | 80 | NS | Platform only | CARAT smartphone app for asthma monitoring with e-mail feedback | R | 6 mo | No difference between groups for medication adherence | No significant difference between groups | — |
| 92 | Adolescent or younger | 40 | NS | Platform only | CHANGE Asthma smartphone app with education and AMP | R | 4 mo | — | No significant difference in cACT between groups | — |
| 93 | Adolescent or younger | 234 | NS | Platform only | ADAPT interactive smartphone app including CARAT | R | 6 mo | Significant improvement in self-reported adherence patients with poor baseline adherence with or without uncontrolled asthma; no significant difference overall | No significant difference in CARAT score between groups | — |

(continued)

TABLE V. (Continued)

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|-----------------------|-----|-----------------|-------------------|--------------------------------------------------------------------------|------------|--------------|--------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------|
| 94 | Adolescent or younger | 87 | NS | Platform only | ADAPT interactive smartphone app including CARAT | R | 6 mo | No significant change in self-reported adherence | Change in CARAT score during therapy not reported | — |
| 95 | Adolescent or younger | 15 | NS | Platform only | mASMAA interactive SMS asthma diary system | SA | 2 wk | Anecdotal reports of improved adherence | Anecdotal reports of improved control over asthma symptoms | — |
| 96 | Adults only | 107 | NS | Platform only | Web platform for asthma monitoring, AMP and HCP contact | R | 30 mo | — | Significant and sustained (≥ 30 mo) improvement in asthma control via AQLQ and ACQ | — |
| 97 | Adolescent or younger | 24 | All | Platform only | Web-based peak flow and symptom tracker with feedback | SA | 2-15 mo | — | Significant reduction in number of wheezing episodes | Significant reduction in doctor or clinic visits |
| 98 | Adolescent or younger | 90 | All | Platform only | Web-based peak flow and symptom tracker with education, AMP and feedback | R | 1 y | No significant difference in self-reported medication adherence between groups | Significant improvement in ACQ scores but not sustained beyond 12 mo | — |
| 99 | Adults only | 200 | Moderate | Platform only | Web-based peak flow and symptom tracker with AMP and feedback | R | 1 y | — | Significant improvement in ACQ for patients with partly controlled or uncontrolled asthma only | — |
| 100 | Adults only | 106 | NS | Platform only | Web-based AMP with SMS feedback | R | 12 mo | Change in adherence during therapy not reported | No significant difference in ACT between groups | No significant difference in hospitalizations or ED visits between groups |
| 101 | Adolescent or younger | 24 | NS | Platform only | AsthmaCare smartphone asthma management app | SA | 30 d | Increased adherence in 85% of subjects | — | — |
| 102 | Adults only | 22 | NS | Platform only | Web-based AMP and symptom/peak flow diary with feedback | SA | 8 d | Change in adherence during therapy not reported | Significant improvement in CACG symptom benchmarks | No significant difference in urgent health care visits |
| 103 | Adolescents | 422 | All | Platform only | Web-based platform with tailored feedback to questions | R | 12 mo | — | Significant reduction in symptom days | No difference in frequency of ED visits |

| | | | | | | | | | | |
|-----|-----------------------|-----|--------------------|---------|------------------------------------------------------------------|----|------------|---------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------|
| 104 | Adults only | 51 | Moderate to severe | Telemed | EM by phone plus static education | SA | 9 mo | 31.4% of 35 subjects indicated improved adherence | Change in ACT statistically significant, but less than MID | No difference in ED visits or hospitalizations between baseline and end of study |
| 105 | Adolescent or younger | 400 | NS | Telemed | Telemed visits | R | 1 school y | — | Significant improvement in symptom-free days per 2 wk | Reduced likelihood of ED visit or hospitalization for asthma |
| 106 | Adolescent or younger | 210 | NS | Telemed | Online virtual asthma clinic | R | 16 mo | — | Significant improvement in cACT but not ACT, in children and adolescents, respectively | No significant difference between groups |
| 107 | Adults only | 100 | NS | Telemed | Telemed and SMS asthma monitoring | R | 12 mo | — | Statistically, but not clinically, significant increase in ACT score for intervention and control (between-group interaction not reported) | No hospitalizations during study |
| 108 | All ages | 20 | NS | Telemed | Telemed education | SA | 1 y | — | Increase in proportion of well-controlled patients by ACT score | — |
| 109 | Adolescent or younger | 210 | NS | Telemed | Telemed visits | R | 16 mo | — | Significant increase in cACT but no difference in ACT for teenagers, after 16 mo | Online asthma management was associated with fewer outpatient clinic visits |
| 110 | Adolescent or younger | 50 | Severe | Telemed | Telemonitoring of FEV ₁ and intervention if necessary | R | 12 mo | — | No significant difference between groups for exacerbations | — |
| 111 | Adolescent or younger | 48 | NS | Telemed | Telemed evaluation compared with face-to-face | SA | 7 mo | — | — | — |
| 112 | Adults only | 422 | NS | Telemed | Telemed EM | R | 2 y | — | Significantly improved ACT scores after 2 y | Significant reduction in asthma-related hospitalization |
| 113 | Adolescent or younger | 362 | NS | Telemed | Telemed EM | R | 1 y | — | Significant reduction in proportion of children with poorly controlled asthma | Urgent care events decreased in both groups with no significant difference between groups |

(continued)

TABLE V. (Continued)

| Reference | Patient age group | n | Asthma severity | Intervention type | Intervention details | Study type | Study period | Effect on ICS adherence with intervention | Effect on asthma impairment or symptoms with intervention | Effect on health care use and/or hospitalization with intervention |
|-----------|-----------------------|-----|-----------------|-------------------|-----------------------------------------|------------|--------------|---------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------|
| 114 | Adolescent or younger | 393 | NS | Telemed | Telemed EM | R | 6 mo | Significant increase in caregiver-reported adherence but no significant difference in asthma medication ratio | No statistically significant difference between groups in reported symptom-free days | — |
| 115 | Adults only | 30 | NS | Telemed | Telemed education and inhaler technique | SA | 2 wk | — | — | — |
| 116 | Adolescent or younger | 295 | NS | Telemed | Telemed education | R | 1 y | — | — | Significant reduction in ED visits and hospitalizations during follow-up |
| 117 | Adults only | 98 | NS | Telemed | Telemed education | R | 3 mo | Within-group improvement in adherence in intervention group | No significant between-group difference in ACT; within-group improvement in intervention group | — |

AMP, asthma management plan; (c)ACT, (childhood) Asthma Control Test; ACQ, Asthma Control Questionnaire; AQLQ, Asthma Quality of Life Questionnaire; EM, education or motivation; EMA, ecological momentary association; FEV₁, forced expiratory volume; HCP, health care professional; ICS, inhaled corticosteroid; MID, minimal important difference; PEFR, peak expiratory flow rate; PEV, peak expiratory volume; SA, single-arm; NS, not specified; R, randomized; SABA, short-acting β-agonist; SMS, short message service; Telemed, telemedicine intervention.

improve individual asthma outcomes through treatment personalization and daily disease management. Data analysis may also assist the understanding of relations and causal mechanisms among patient characteristics, environment, and asthma symptoms and exacerbations, ultimately leading to the development of patient digital phenotypes.

However, beyond the challenge of developing suitable technologies, considerable financial, technological, and political barriers still need to be overcome. The development of artificial intelligence protocols to interpret and apply the huge amount of monitoring data involved will be important. There will also be a considerable emphasis on the infrastructure necessary to incorporate continuous monitoring into patients' overall health care to support physicians. Such a fundamental change in the organization and delivery of health care will also have significant effects on liability for physicians making artificial intelligence-augmented decisions.^{133,134} Large-scale personal health data collection naturally generates concern regarding data security and use, and patients and physicians must be confident about how their data will be used, who will have access to it, and the extent to which individual patients will own their data.¹³⁵

Regardless of pharmacological therapy, consultations can already be implemented regularly by videoconference, which is especially important for patients with reduced mobility or reduced free time, or in remote locations; videoconferencing could also be used to provide support between face-to-face appointments. In the context of a pandemic such as COVID-19, reducing face-to-face contact between clinicians and selected patients has been a necessity. The use of telemedicine and video consultation was implemented as an elegant solution to this challenge for patient assessment and education, both of which can be delivered remotely.^{136,137} One related effect of the current pandemic situation is a large-scale uptake of videoconference technology for work and social contact in the general population, which could help acclimatize patients and physicians to accessing and providing health care in this way. We consider it likely that many of the benefits of telemedicine realized during the pandemic, initially through necessity, will continue to gain increasing acceptance and form an integral part of respiratory care in the future.^{137,138}

Limitations

Conclusions from this scoping review are limited by heterogeneity in terms of the wide variety of study designs and outcome measures and the different ways of implementing similar interventions for patient-independent and patient-dependent initiatives. In addition, study heterogeneity limited our interpretation of different outcomes for the same intervention category (eg, the changes observed in adherence were not necessarily measured in the same studies as changes in ACT). The ACT was the most common measure of asthma burden and is a subjective measure of patients' perception of risk for an asthma exacerbation. Changes in asthma exacerbations were not commonly used as a primary outcome measure, possibly because the expected number of exacerbations in study populations was not sufficiently frequent to make this a useful end point.

Because patients were generally monitored for no more than 12 months (and commonly only 3 or 6 months), the long-term effects of the various interventions are unknown. Studies also

typically did not account for seasonality or incorporate environmental information (such as aeroallergen or air pollution levels). Most of the studies analyzed did not specify participants' asthma severity at baseline and lacked stratification by asthma severity. An abundance of studies also reported outcomes only at single specific time points or as a single assessment for the whole study period, rather than as change over time associated with a specific intervention.

The relation between asthma controller adherence and health care use is an important issue, but one that is challenging to study. In addition to the acknowledged problems with the accuracy of self-reported adherence,^{48,139,140} in a general population of treated asthma patients who are not under close clinical supervision, it would be expected that the frequency of emergency or unplanned clinical consultation would be low, and therefore statistically significant changes would be difficult to detect. The studies identified for this review rarely assessed health care use systematically, and few reported on both adherence and health care use. Furthermore, correlating changes in adherence with changes in use outcomes is confounded by several patient-related and sociobehavioral factors including symptom severity, ethnicity, level of educational achievement, household income, and attitude toward treatment.¹⁴¹

Therefore, the fact that reductions in health care use were not reported with digital asthma interventions in the studies discussed in this review does not mean that these interventions are not associated with benefit in terms of health care resource use in the real world. The patient populations, intervention assessments, and data collection methods might not have been capable of detecting health care use changes accurately, even where these may have occurred. Further research into the effects of adherence on health care use is undoubtedly needed, and considerable effort will be required to account for these factors in the design of future studies assessing digital interventions. Furthermore, future research into digital technology as a part of asthma interventions would benefit from a convention on standard approaches to outcomes and their measurement to enhance similarity in the ways in which we assess asthma treatment.

CONCLUSIONS

Digital health interventions are acceptable to asthma patients and clinicians, and the relative ubiquity of mobile Internet- and Bluetooth-connected devices in everyday life makes real-time monitoring of asthma treatment and symptoms extremely feasible. However, an effective digital intervention must fulfill several important design criteria. Both inhaler devices and software should be intuitive and easy to use, and hardware should be unobtrusive with accurate and objective measurement of adherence and other parameters, and be seamlessly integrated with software providing convenient assessment of symptoms and asthma control. Software should be designed to educate and maintain patients' engagement (at a variety of different ages), provide clear communication of hardware- and software-measured parameters for access by patients and clinicians, and be customizable by patients to fit into their lives and routines. Motivating patients to take an active role in managing their condition and maintaining their interest in doing so is of

paramount importance in designing a successful future digital intervention.

Acknowledgments

Medical writing support for the development of this article was provided by Ian C. Grieve, PhD, of Ashfield Healthcare Communications, part of UDG Healthcare plc, and was funded by Teva Pharmaceuticals.

REFERENCES

1. The Global Asthma Report 2018. Auckland, New Zealand: Global Asthma Network; 2018.
2. Levy ML. The national review of asthma deaths: what did we learn and what needs to change? *Breathe (Sheff)* 2015;11:14-24.
3. Pinnock H, Parke HL, Panagioti M, Daines L, Pearce G, Epiphaniou E, et al. Systematic meta-review of supported self-management for asthma: a health-care perspective. *BMC Med* 2017;15:64.
4. Gruffydd-Jones K. Unmet needs in asthma. *Ther Clin Risk Manag* 2019;15:409-21.
5. Akinbami LJ, Salo PM, Cloutier MM, Wilkerson JC, Elward KS, Mazurek JM, et al. Primary care clinician adherence with Asthma Guidelines: the National Asthma Survey of Physicians. *J Asthma* 2020;57:543-55.
6. Cloutier MM, Salo PM, Akinbami LJ, Cohn RD, Wilkerson JC, Diette GB, et al. Clinician agreement, self-efficacy, and adherence with the guidelines for the diagnosis and management of asthma. *J Allergy Clin Immunol Pract* 2018; 6:886-894.e4.
7. Morrison D, Mair FS, Yardley L, Kirby S, Thomas M. Living with asthma and chronic obstructive airways disease: using technology to support self-management – an overview. *Chron Respir Dis* 2017;14:407-19.
8. Mann DM, Chen J, Chunara R, Testa PA, Nov O. COVID-19 transforms health care through telemedicine: evidence from the field. *J Am Med Inform Assoc* 2020;27:1132-5.
9. Pham MT, Rajić A, Grieg JD, Sargeant JM, Papadopoulos A, McEwan SA. A scoping review of scoping reviews: advancing the approach and enhancing the consistency. *Res Synth Methods* 2014;5:371-85.
10. Tricco AC, Lillie E, Zarin W, O'Brien KK, Colquhoun H, Levac D, et al. PRISMA extension for scoping reviews (PRISMA-ScR): checklist and explanation. *Ann Intern Med* 2018;169:467-73.
11. Chan AHY, Stewart AW, Harrison J, Black PN, Mitchell EA, Foster JM. Electronic adherence monitoring device performance and patient acceptability: a randomized control trial. *Expert Rev Med Devices* 2017;14:401-11.
12. Chan AH, Stewart AW, Harrison J, Camargo CA, Black PN, Mitchell EA. The effect of an electronic monitoring device with audiovisual reminder function on adherence to inhaled corticosteroids and school attendance in children with asthma: a randomised controlled trial. *Lancet Respir Med* 2015;3:210-9.
13. Cvietusa PJ, Goodrich GK, Shoup JA, Steffen DA, Tacinas C, Wagner NM, et al. Implementing health care technology research into practice to improve adult asthma management. *J Allergy Clin Immunol Pract* 2019;7:908-14.
14. Bender BG, Cvietusa PJ, Goodrich GK, Lowe R, Nuanes HA, Rand C, et al. Pragmatic trial of health care technologies to improve adherence to pediatric asthma treatment: a randomized clinical trial. *JAMA Pediatr* 2015; 169:317-23.
15. Lee SH, Song WJ, Park HW, Kang MG, Kim SH, Park HK, et al. Multifaceted interventions to reduce acute exacerbations in elderly asthmatics. *Asia Pac Allergy* 2018;8:e1.
16. Britto MT, Rohan JM, Dodds CM, Byczkowski TL. A randomized trial of user-controlled text messaging to improve asthma outcomes: a pilot study. *Clin Pediatr (Phila)* 2017;56:1336-44.
17. Johnson KB, Patterson BL, Ho YX, Chen Q, Nian H, Davison CL, et al. The feasibility of text reminders to improve medication adherence in adolescents with asthma. *J Am Med Inform Assoc* 2016;23:449-55.
18. Strandbygaard U, Thomsen SF, Backer V. A daily SMS reminder increases adherence to asthma treatment: a three-month follow-up study. *Respir Med* 2010;104:166-71.

19. Britto MT, Munafo JK, Schoettler PJ, Vockell AL, Wimberg JA, Yi MS. Pilot and feasibility test of adolescent-controlled text messaging reminders. *Clin Pediatr (Phila)* 2012;51:114-21.
20. Vollmer WM, Feldstein A, Smith DH, Dubanoski JP, Waterbury A, Schneider JL, et al. Use of health information technology to improve medication adherence. *Am J Manag Care* 2011;17:SP79-87.
21. Sleath B, Carpenter DM, Davis SA, Watson CH, Lee C, Loughlin CE, et al. Improving youth question-asking and provider education during pediatric asthma visits. *Patient Educ Couns* 2018;101:1051-7.
22. Thomas M, Bruton A, Little P, Holgate S, Lee A, Yardley L, et al. A randomised controlled study of the effectiveness of breathing retraining exercises taught by a physiotherapist either by instructional DVD or in face-to-face sessions in the management of asthma in adults. *Health Technol Assess* 2017;21:1-162.
23. Lv Y, Zhao H, Liang Z, Dong H, Liu L, Zhang D, et al. A mobile phone short message service improves perceived control of asthma: a randomized controlled trial. *Telemed J E Health* 2012;18:420-6.
24. Cowden JD, Wilkerson-Amendell S, Weathers L, Gonzalez ED, Dinakar C, Westbrook DH, et al. The talking card: randomized controlled trial of a novel audio-recording tool for asthma control. *Allergy Asthma Proc* 2015;36:e86-91.
25. Lau AY, Arguel A, Dennis S, Liaw ST, Coiera E. Why didn't it work? Lessons from a randomized controlled trial of a Web-based personally controlled health management system for adults with asthma. *J Med Internet Res* 2015;17:e283.
26. Kim YH, Yoo KH, Yoo JH, Kim TE, Kim DK, Park YB, et al. The need for a well-organized, video-assisted asthma education program at Korean primary care clinics. *Tuberc Respir Dis (Seoul)* 2017;80(2):169-78.
27. Lipszyc JC, Gotzev S, Scarborough J, Liss GM, Gupta S, Tarlo SM. Evaluation of the efficacy of a web-based work-related asthma educational tool. *J Asthma* 2016;53:1071-5.
28. Hughes M, Murphy M. Evaluation of a pilot national online asthma e-learning program for secondary school students. *Issues Compr Pediatr Nurs* 2014;37:136-46.
29. Burns P, Jones SC, Iverson D, Caputi P. AsthmaWise — a field of dreams? The results of an online education program targeting older adults with asthma. *J Asthma* 2013;50:737-44.
30. Kercsmar CM, Sorkness CA, Calatroni A, Gergen PJ, Bloomberg GR, Gruchalla RS, et al. A computerized decision support tool to implement asthma guidelines for children and adolescents. *J Allergy Clin Immunol* 2019;143:1760-8.
31. Kenyon CC, Gruschow SM, Quarshie WO, Griffis H, Leach MC, Zorc JJ, et al. Controller adherence following hospital discharge in high risk children: a pilot randomized trial of text message reminders. *J Asthma* 2019;56:95-103.
32. Sulaiman I, Seheult J, MacHale E, Boland F, O'Dwyer SM, Rapcan V, et al. A method to calculate adherence to inhaled therapy that reflects the changes in clinical features of asthma. *Ann Am Thorac Soc* 2016;13:1894-903.
33. Killane I, Sulaiman I, MacHale E, Breathnach A, Taylor TE, Holmes MS, et al. Predicting asthma exacerbations employing remotely monitored adherence. *Health Technol Lett* 2016;3:51-5.
34. Hernández CR, Fernández MN, Sanmartín AP, Roibas CM, Domínguez LC, Rial MIB, et al. AF validation of the portable Air-Smart Spirometer. *PLoS One* 2018;13:e0192789.
35. Gahleitner F, Legg J, Holland E, Pearson S, Roberts G. The validity and acceptability of a text-based monitoring system for pediatric asthma studies. *Pediatr Pulmonol* 2016;51:5-12.
36. Patel M, Perrin K, Pritchard A, Williams M, Wijesinghe M, Weatherall M, et al. Accuracy of patient self-report as a measure of inhaled asthma medication use. *Respirology* 2013;18:546-52.
37. Vasbinder E, Dahhan N, Wolf B, Zoer J, Blankman E, Bosman D, et al. The association of ethnicity with electronically measured adherence to inhaled corticosteroids in children. *Eur J Clin Pharmacol* 2013;69:683-90.
38. Sulaiman I, Seheult J, MacHale E, D'Arcy S, Boland F, McCrory K, et al. Irregular and ineffective: a quantitative observational study of the time and technique of inhaler use. *J Allergy Clin Immunol Pract* 2016;4:900-909.e2.
39. Beerhuizen T, Voorend-van Bergen S, van den Hout WB, Vaessen-Verberne AA, Brackel HJ, Landstra AM, et al. Cost-effectiveness of FENO-based and web-based monitoring in paediatric asthma management: a randomised controlled trial. *Thorax* 2016;71:607-13.
40. Voorend-van Bergen S, Vaessen-Verberne AA, Brackel HJ, Landstra AM, van den Berg NJ, Hop WC, et al. Pijnburg Monitoring strategies in children with asthma: a randomised controlled trial. *Thorax* 2015;70:543-50.
41. Rhee H, Belyea MJ, Sterling M, Bocko MF. Evaluating the validity of an automated device for asthma monitoring for adolescents: correlational design. *J Med Internet Res* 2015;17:e234.
42. Mulvaney SA, Ho YX, Cala CM, Chen Q, Nian H, Patterson BL, et al. Assessing adolescent asthma symptoms and adherence using mobile phones. *J Med Internet Res* 2013;15:e141.
43. Uysal MA, Mungan D, Yorgancioglu A, Yildiz F, Akgun M, Gemicioğlu B, et al. Asthma control test via text messaging: could it be a tool for evaluating asthma control? *J Asthma* 2013;50:1083-9.
44. MacDonell K, Gibson-Scipio W, Lam P, Naar-King S, Chen X. Text messaging to measure asthma medication use and symptoms in urban African American emerging adults: a feasibility study. *J Asthma* 2012;49:1092-6.
45. Voorend-van Bergen S, Vaessen-Verberne AA, Landstra AM, Brackel HJ, van den Berg NJ, Caudri D, et al. Monitoring childhood asthma: web-based diaries and the asthma control test. *J Allergy Clin Immunol* 2014;133:1599-1605.e2.
46. Montalbano L, Ferrante G, Cilluffo G, Gentile M, Arrigo M, La Guardia D, et al. Targeting quality of life in asthmatic children: the MyTEP pilot randomized trial. *Respir Med* 2019;153:14-9.
47. Chan YY, Wang P, Rogers L, Tignor N, Zweig M, Hershman SG, et al. The Asthma Mobile Health Study, a large-scale clinical observational study using ResearchKit. *Nat Biotechnol* 2017;35:354-62.
48. Mosnaim G, Li H, Martin M, Richardson D, Belice PJ, Avery E, et al. The impact of peer support and mp3 messaging on adherence to inhaled corticosteroids in minority adolescents with asthma: a randomized, controlled trial. *J Allergy Clin Immunol Pract* 2013;1:485-93.
49. Petrie KJ, Perry K, Broadbent E, Weinman J. A text message programme designed to modify patients' illness and treatment beliefs improves self-reported adherence to asthma preventer medication. *Br J Health Psychol* 2012;17:74-84.
50. Klok T, Kaptein AA, Duiverman EJ, Brand PL. It's the adherence, stupid (that determines asthma control in preschool children)! *Eur Respir J* 2014;43:783-91.
51. Bender BG, Apter A, Bogen DK, Dickinson P, Fisher L, Wamboldt FS, et al. Test of an interactive voice response intervention to improve adherence to controller medications in adults with asthma. *J Am Board Fam Med* 2010;23:159-65.
52. Jaimini U, Thirunarayan K, Kalra M, Venkataraman R, Kadariya D, Sheth A. "How is my child's asthma?" Digital phenotype and actionable insights for pediatric asthma. *JMIR Pediatr Parent* 2018;1:e11988.
53. Cao Y, Lin S-H, Zhu D, Xu F, Chen Z-H, Shen H-H, et al. WeChat public account use improves clinical control of cough-variant asthma: a randomized controlled trial. *Med Sci Monit* 2018;24:1524-32.
54. Mikalsen IB, Nassehi D, Øymar K. Vortex whistle and smart phone application for peak flow recordings in asthmatic children: a feasibility study. *Telemed J E Health* 2019;25:1077-82.
55. Zairina E, Abramson MJ, McDonald CF, Li J, Dharmasiri T, Stewart K, et al. Telehealth to improve asthma control in pregnancy: a randomized controlled trial. *Respirology* 2016;21:867-74.
56. Speck AL, Hess M, Baptist AP. An electronic asthma self-management intervention for young African American adults. *J Allergy Clin Immunol Pract* 2016;4:89-95.e2.
57. Morrison D, Wyke S, Saunderson K, McConnachie A, Agur K, Chaudhuri R, et al. Findings from a pilot Randomised trial of an Asthma Internet Self-management Intervention (RAISIN). *BMJ Open* 2016;6:e009254.
58. Weinstein AG, Singh A, Laurenceau JP, Skoner DP, Maiolo J, Sharara R, et al. A pilot study of the effect of an educational Web application on asthma control and medication adherence. *J Allergy Clin Immunol Pract* 2019;7:1497-506.
59. Merchant RK, Inamdar R, Quade RC. Effectiveness of population health management using the Propeller Health Asthma Platform: a randomized clinical trial. *J Allergy Clin Immunol Pract* 2016;4:455-63.
60. Morton RW, Elphick HE, Rigby AS, Daw WJ, King DA, Smith LJ, et al. STAAR: a randomised controlled trial of electronic adherence monitoring with reminder alarms and feedback to improve clinical outcomes for children with asthma. *Thorax* 2017;72:347-54.
61. Kenyon CC, Chang J, Wynter SA, Fowler JC, Long J, Bryant-Stephens TC. Electronic adherence monitoring in a high-utilizing pediatric asthma cohort: a feasibility study. *JMIR Res Protoc* 2016;5:e132.
62. Cushing A, Manice MP, Ting A, Parides MK. Feasibility of a novel mHealth management system to capture and improve medication adherence among adolescents with asthma. *Patient Prefer Adherence* 2016;10:2271-5.
63. Merchant R, Szefer SJ, Bender BG, Tuffli M, Barrett MA, Gondalia R, et al. Impact of a digital health intervention on asthma resource utilization. *World Allergy Organ J* 2018;11:28.
64. Zhou Y, Lu Y, Zhu H, Zhang Y, Li Y, Yu Q. Short-term effect of a smart nebulizing device on adherence to inhaled corticosteroid therapy in Asthma Predictive Index-positive wheezing children. *Patient Prefer Adherence* 2018;12:861-8.

65. Grossman B, Conner S, Mosnaim G, Albers J, Leigh J, Jones S, et al. Application of Human Augmentics: a persuasive asthma inhaler. *J Biomed Inform* 2017;67:51-8.
66. Van Sickle D, Magzamen S, Truelove S, Morrison T. Remote monitoring of inhaled bronchodilator use and weekly feedback about asthma management: an open-group, short-term pilot study of the impact on asthma control. *PLoS One* 2013;8:e55335.
67. Barrett MA, Humblet O, Marcus JE, Henderson K, Smith T, Eid N, et al. Effect of a mobile health, sensor-driven asthma management platform on asthma control. *Ann Allergy Asthma Immunol* 2017;119:415-21.
68. Mosnaim G, Li H, Martin M, Richardson D, Belice PJ, Avery E, et al. A tailored mobile health intervention to improve adherence and asthma control in minority adolescents. *J Allergy Clin Immunol Pract* 2015;3:288-290.e1.
69. Foster JM, Usherwood T, Smith L, Sawyer SM, Xuan W, Rand CS, et al. Inhaler reminders improve adherence with controller treatment in primary care patients with asthma. *J Allergy Clin Immunol* 2014;134:1260-1268.e3.
70. Kuipers E, Poot CC, Wensing M, Chavannes NH, de Smet PA, Teichert M. Self-Management Maintenance Inhalation Therapy With eHealth (SELFIE): observational study on the use of an electronic monitoring device in respiratory patient care and research. *J Med Internet Res* 2019;21:e13551.
71. Barrett M, Combs V, Su JG, Henderson K, Tuffli M, Simrall G, et al. AIR Louisville: addressing asthma with technology, crowdsourcing, cross-sector collaboration, and policy. *Health Affairs* 2018;37:525-34.
72. Morita PP, Yeung MS, Ferrone M, Taite AK, Madeley C, Stevens Lavigne A, et al. A patient-centered mobile health system that supports asthma self-management (breathe): design, development, and utilization. *JMIR Mhealth Uhealth* 2019;7:e10956.
73. Pool AC, Kraschnewski JL, Poger JM, Smyth J, Stuckey HL, Craig TJ, et al. Impact of online patient reminders to improve asthma care: a randomized controlled trial. *PLoS One* 2017;12:e0170447.
74. Cook KA, Modena BD, Simon RA. Improvement in asthma control using a minimally burdensome and proactive smartphone application. *J Allergy Clin Immunol Pract* 2016;4:730-737.e1.
75. Ahmed S, Ernst P, Bartlett SJ, Valois MF, Zaihra T, Paré G, et al. The effectiveness of Web-based asthma self-management system, My Asthma Portal (MAP): a pilot randomized controlled trial. *J Med Internet Res* 2016;18:e313.
76. Cingi C, Yorgancioglu A, Cingi CC, Oguzulgen K, Muluk NB, Ulusoy S, et al. The "physician on call patient engagement trial" (POPET): measuring the impact of a mobile patient engagement application on health outcomes and quality of life in allergic rhinitis and asthma patients. *Int Forum Allergy Rhinol* 2015;5:487-97.
77. Lv S, Ye X, Wang Z, Xia W, Qi Y, Wang W, et al. A randomized controlled trial of a mobile application-assisted nurse-led model used to improve treatment outcomes in children with asthma. *J Adv Nurs* 2019;75:3058-67.
78. Stukus DR, Farooqui N, Strothman K, Ryan K, Zhao S, Stevens JH, et al. Real-world evaluation of a mobile health application in children with asthma. *Ann Allergy Asthma Immunol* 2018;120:395-400.e1.
79. Perry TT, Marshall A, Berlinski A, Rettiganti M, Brown RH, Randle SM, et al. Smartphone-based vs paper-based asthma action plans for adolescents. *Ann Allergy Asthma Immunol* 2017;118:298-303.
80. Koufopoulos JT, Conner MT, Gardner PH, Kellar I. A Web-based and mobile health social support intervention to promote adherence to inhaled asthma medications: randomized controlled trial. *J Med Internet Res* 2016;18:e122.
81. Fiks AG, DuRivage N, Mayne SL, Finch S, Ross ME, Giacomini K, et al. Adoption of a portal for the primary care management of pediatric asthma: a mixed-methods implementation study. *J Med Internet Res* 2016;18:e172.
82. Kim MY, Lee SY, Jo EJ, Lee SE, Kang MG, Song WJ, et al. Feasibility of a smartphone application based action plan and monitoring in asthma. *Asia Pac Allergy* 2016;6:174-80.
83. Kolmodin MacDonell K, Naar S, Gibson-Scipio W, Lam P, Secord E. The Detroit Young Adult Asthma Project: pilot of a technology-based medication adherence intervention for African-American emerging adults. *J Adolesc Health* 2016;59:465-71.
84. Wiecha JM, Adams WG, Rybin D, Rizzodepaoli M, Keller J, Clay JM. Evaluation of a web-based asthma self-management system: a randomized controlled pilot trial. *BMC Pulm Med* 2015;15:17.
85. Burbank AJ, Lewis SD, Hewes M, Schellhase DE, Rettiganti M, Hall-Barrow J, et al. Mobile-based asthma action plans for adolescents. *J Asthma* 2015;52:583-6.
86. Araújo L, Jacinto T, Moreira A, Castel-Branco MG, Delgado L, Costa-Pereira A. Clinical efficacy of web-based versus standard asthma self-management. *J Investig Allergol Clin Immunol* 2012;22:28-34.
87. Seid M, D'Amico EJ, Varni JW, Munafò JK, Britto MT, Kercsmar CM, et al. The in vivo adherence intervention for at risk adolescents with asthma: report of a randomized pilot trial. *J Pediatr Psychol* 2012;37:390-403.
88. Liu WT, Huang CD, Wang CH, Lee KY, Lin SM, Kuo HP. A mobile telephone-based interactive self-care system improves asthma control. *Eur Respir J* 2011;37:310-7.
89. Prabhakaran L, Chee WY, Chua KC, Abisheganaden J, Wong WM. The use of text messaging to improve asthma control: a pilot study using the mobile phone short messaging service (SMS). *J Telemed Telecare* 2010;16:286-90.
90. Lathy P, Neo LP, Gan CC, Tham LM, Ng C, Yap CW, et al. Effectiveness of the ecare programme: a short message service (SMS) for asthma monitoring. *Ann Acad Med Singap* 2018;47:233-6.
91. Kuipers E, Wensing M, de Smet P, Teichert M. Self-management research of asthma and good drug use (SMARAGD study): a pilot trial. *Int J Clin Pharm* 2017;39:888-96.
92. Real FJ, Beck AF, DeBlasio D, Zackoff M, Henize A, Xu Y, et al. Dose Matters: a smartphone application to improve asthma control among patients at an urban pediatric primary care clinic. *Games Health J* 2019;8:357-65.
93. Kosse RC, Bouvy ML, de Vries TW, Koster ES. Effect of a mHealth intervention on adherence in adolescents with asthma: a randomized controlled trial. *Respir Med* 2019;149:45-51.
94. Kosse RC, Bouvy ML, Belitser SV, de Vries TW, van der Wal PS, Koster ES. Effective engagement of adolescent asthma patients with mobile health-supporting medication adherence. *JMIR Mhealth Uhealth* 2019;7:e12411.
95. Rhee H, Miner S, Sterling M, Halterman JS, Fairbanks E. The development of an automated device for asthma monitoring for adolescents: methodologic approach and user acceptability. *JMIR Mhealth Uhealth* 2014;2:e27.
96. van Gaalen JL, Beerthuis T, van der Meer V, van Reisen P, Redelijkheid GW, Snoeck-Stroband JB, et al. Long-term outcomes of internet-based self-management support in adults with asthma: randomized controlled trial. *J Med Internet Res* 2013;15:e188.
97. Arnold RJ, Stingone JA, Claudio L. Computer-assisted school-based asthma management: a pilot study. *JMIR Res Protoc* 2012;1:e15.
98. Rikkers-Mutsaerts ER, Winters AE, Bakker MJ, van Stel HF, van der Meer V, de Jongste JC, et al. Internet-based self-management compared with usual care in adolescents with asthma: a randomized controlled trial. *Pediatr Pulmonol* 2012;47:1170-9.
99. van der Meer V, van Stel HF, Bakker MJ, Roldaan AC, Assendelft WJ, Sterk PJ, et al. Weekly self-monitoring and treatment adjustment benefit patients with partly controlled and uncontrolled asthma: an analysis of the SMASHING study. *Respir Res* 2010;11:74.
100. Poureslami I, Shum J, Lester RT, Tavakoli H, Dorscheid DR, FitzGerald JM. A pilot randomized controlled trial on the impact of text messaging check-ins and a web-based asthma action plan versus a written action plan on asthma exacerbations. *J Asthma* 2018;16:1-13.
101. Farooqui N, Phillips G, Barrett C, Stukus D. Acceptability of an interactive asthma management mobile health application for children and adolescents. *Ann Allergy Asthma Immunol* 2015;114:527-9.
102. Licskai C, Sands TW, Ferrone M. Development and pilot testing of a mobile health solution for asthma self-management: asthma action plan smartphone application pilot study. *Can Respir J* 2013;20:301-6.
103. Joseph CL, Ownby DR, Havstad SL, Saltzgaber J, Considine S, Johnson D, et al. Evaluation of a web-based asthma management intervention program for urban teenagers: reaching the hard to reach. *J Adolesc Health* 2013;52:419-26.
104. Rasulnia M, Burton BS, Ginter RP, Wang TY, Pleasants RA, Green CL, et al. Assessing the impact of a remote digital coaching engagement program on patient-reported outcomes in asthma. *J Asthma* 2018;55:795-800.
105. Halterman JS, Fagnano M, Tajon RS, Tremblay P, Wang H, Butz A, et al. Effect of the School-Based Telemedicine Enhanced Asthma Management (SB-TEAM) program on asthma morbidity: a randomized clinical trial. *JAMA Pediatr* 2018;172:e174938.
106. van den Wijngaart LS, Roukema J, Boehmer ALM, Brouwer ML, Hugen CAC, Niers LEM, et al. A virtual asthma clinic for children: fewer routine outpatient visits, same asthma control. *Eur Respir J* 2017;50:1700471.
107. Nemanic T, Sarc I, Skrgat S, Flezar M, Cukjati I, Marc Malovrh M. Telemonitoring in asthma control: a randomized controlled trial. *J Asthma* 2019;56:782-90.
108. Brown W, Scott D, Friesner D, Schmitz T. Impact of telepharmacy services as a way to increase access to asthma care. *J Asthma* 2017;54:961-7.
109. Van Den Wijngaart LS, Kievit W, Roukema J, Boehmer AL, Brouwer ML, Hugen CA, et al. The virtual asthma clinic for children: a cost-effectiveness analysis. *Pediatr Pulmonol* 2016;51:S65.

110. Deschilde A, Béghin L, Salleron J, Iliescu C, Thumerelle C, Santos C, et al. Home telemonitoring (forced expiratory volume in 1 s) in children with severe asthma does not reduce exacerbations. *Eur Respir J* 2012;39:290-6.
111. Gattu R, Scollan J, DeSouza A, Devereaux D, Weaver H, Agthe AG. Telemedicine: a reliable tool to assess the severity of respiratory distress in children. *Hosp Pediatr* 2016;6:476-82.
112. Patel MR, Song PXX, Sanders G, Nelson B, Kaltsas E, Thomas LJ, et al. A randomized clinical trial of a culturally responsive intervention for African American women with asthma. *Ann Allergy Asthma Immunol* 2017;118:212-9.
113. Garbutt JM, Banister C, Highstein G, Sterkel R, Epstein J, Bruns J, et al. Telephone coaching for parents of children with asthma: Impact and lessons learned. *Arch Pediatr Adolesc Med* 2010;164:625-30.
114. Perry TT, Halterman JS, Brown RH, Luo C, Randle SM, Hunter CR, et al. Results of an asthma education program delivered via telemedicine in rural schools. *Ann Allergy Asthma Immunol* 2018;120:401-8.
115. Nelson P, Young HN, Knobloch MJ, Griesbach SA. Telephonic monitoring and optimization of inhaler technique. *Telemed J E Health* 2011;17:734-40.
116. Coughney K, Klein G, West C, Diamond JJ, Santana A, McCarville E, et al. The child asthma link line: a coalition-initiated, telephone-based, care coordination intervention for childhood asthma. *J Asthma* 2010;47:303-9.
117. Young HN, Havican SN, Griesbach S, Thorpe JM, Chewing BA, Sorkness CA. Patient and pharmacist telephonic encounters (PARTE) in an underserved rural patient population with asthma: results of a pilot study. *Telemed J E Health* 2012;18:427-33.
118. McKibben S, De Simoni A, Bush A, Thomas M, Griffiths C. The use of electronic alerts in primary care computer systems to identify the over-prescription of short-acting beta2-agonists in people with asthma: a systematic review. *NPJ Prim Care Resp Med* 2017;27:30.
119. Kosse RC, Bouvy ML, de Vries TW, Koster ES. Evaluation of a mobile health intervention to support asthma self-management and adherence in the pharmacy. *Int J Clin Pharm* 2019;41:452-9.
120. Hollenbach JP, Cushing A, Melvin E, McGowan B, Cloutier MM, Manice M. Understanding clinicians' attitudes toward a mobile health strategy to childhood asthma management: a qualitative study. *J Asthma* 2017;54:754-60.
121. Roberts C, Sage A, Geryk L, Sleath B, Carpenter D. Adolescent preferences and design recommendations for an asthma self-management app: mixed-methods study. *JMIR Form Res* 2018;2:e10055.
122. Sage A, Roberts C, Geryk L, Sleath B, Tate D, Carpenter D. A self-regulation theory-based asthma management mobile app for adolescents: a usability assessment. *JMIR Hum Factors* 2017;4:e5.
123. Elias P, Rajan NO, McArthur K, Dacso CC. InSpire to promote lung assessment in youth: evolving the self-management paradigms of young people with asthma. *Med 2 0* 2013;2:e1.
124. Hui CY, McKinstry B, Walton R, Pinnock H. A mixed method observational study of strategies to promote adoption and usage of an application to support asthma self-management. *J Innov Health Inform* 2019;25:243-53.
125. Doyle R, Albright K, Hurley LP, Chávez C, Stowell M, Dirksen S, et al. Patient perspectives on a text messaging program to support asthma management: a qualitative study. *Health Promot Pract* 2019;20:585-92.
126. Gupta S, Lam Shin Cheung V, Kastner M, Straus S, Kaplan A, Boulet LP, et al. Patient preferences for a touch screen tablet-based asthma questionnaire. *J Asthma* 2019;56:771-81.
127. Merchant R, Inamdar R, Henderson K, Barrett M, Su JG, Riley J, et al. Digital health intervention for asthma: patient-reported value and usability. *JMIR Mhealth Uhealth* 2018;6:e133.
128. McWilliams A, Reeves K, Shade L, Burton E, Tapp H, Courtlandt C, et al. Patient and family engagement in the design of a mobile health solution for pediatric asthma: development and feasibility study. *JMIR Mhealth Uhealth* 2018;6:e68.
129. Foster JM, Reddel HK, Usherwood T, Sawyer SM, Smith L. Patient-perceived acceptability and behaviour change benefits of inhaler reminders and adherence feedback: a qualitative study. *Respir Med* 2017;129:39-45.
130. Roberts CA, Geryk LL, Sage AJ, Sleath BL, Tate DF, Carpenter DM. Adolescent, caregiver, and friend preferences for integrating social support and communication features into an asthma self-management app. *J Asthma* 2016;53:948-54.
131. Burnay E, Cruz-Correia R, Jacinto T, Sousa AS, Fonseca J. Challenges of a mobile application for asthma and allergic rhinitis patient enablement-interface and synchronization. *Telemed J E Health* 2013;19:13-8.
132. Foster J, Reddel HK. Adherence monitoring and inhaler reminders benefit GPs and patients in family practice settings. *Am J Respir Crit Care Med* 2018;201:A4832.
133. Callens S, Galot A, Lamas E. Legal aspects of personal health monitoring. *Stud Health Technol Inform* 2013;187:55-63.
134. Van Biesen W, Decruyenaere J, Sideri K, Cockbain J, Sterckx S. Remote digital monitoring of medication intake: methodological, medical, ethical and legal reflections. *Acta Clin Belg* 2019;23:1-8.
135. Campbell JL, Eyal N, Muslimenta A, Haberer JE. Ethical questions in medical electronic adherence monitoring. *J Gen Intern Med* 2016;31:338-42.
136. Shaker MS, Oppenheimer J, Grayson M, Stukus D, Hartog N, Hsieh EWY, et al. COVID-19: pandemic contingency planning for the allergy and immunology clinic. *J Allergy Clin Immunol Pract* 2020;8:1477-88.
137. Portnoy J, Waller M, Elliott T. Telemedicine in the era of COVID-19. *J Allergy Clin Immunol Pract* 2020;8:1489-91.
138. Portnoy J, Pandya A, Waller M, Elliott T. Telemedicine and emerging technologies for health care in allergy/immunology. *J Allergy Clin Immunol* 2020;145:445-54.
139. Spector SL, Kinsman R, Mawhinney H, Siegel SC, Rachelefsky GS, Katz RM, et al. Compliance of patients with asthma with an experimental aerosolized medication: implications for controlled clinical trials. *J Allergy Clin Immunol* 1986;77:65-70.
140. Kaye L, Gondalia R, Gonzalez C, Adams B, Zhu L, Quan P, et al. Electronic medication monitoring vs self-reported use of inhaled corticosteroids and short-acting beta2-agonists in adult patients with uncontrolled asthma. *J Allergy Clin Immunol* 2020;145:AB59.
141. Apter AJ, Boston RC, George M, Norfleet AL, Tenhave T, Coyne JC, et al. Modifiable barriers to adherence to inhaled steroids among adults with asthma: it's not just black and white. *J Allergy Clin Immunol* 2003;111:1219-26.