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The Role of Mobile Health Technologies in Allergy Care

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The Role of Mobile Health Technologies in Allergy Care: an EAACI Position Paper

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SThe author contributed specific expertise in the area of mobile health/telemedicine in allergology and reviewing the text

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Abstract

Mobile Health (mHealth) uses mobile communication devices such as smartphones and tablet computers to support and improve health-related services, data flow and information, patient selfmanagement, surveillance, and disease management from the moment of first diagnosis to an optimized treatment. The European Academy of Allergy and Clinical Immunology created a task force to assess the state of the art and future potential of mHealth in allergology. The task force endorsed the "Be He@lthy, Be Mobile" WHO initiative and debated the quality, usability, efficiency, advantages, limitations, and risks of mobile solutions for allergic diseases. The results are summarized in this position paper, analyzing also the regulatory background with regard to the "General Data Protection Regulation" and Medical Directives of the European Community. The task force assessed the design, user engagement, content, potential of inducing behavioral change, credibility/accountability, and privacy policies of mHealth products. The perspectives of health care professionals and allergic patients are discussed, underlining the need of thorough investigation for an effective design of mHealth technologies as auxiliary tools to improve quality of care. Within the context of precision medicine, these could facilitate the change in perspective from clinician- to patient-centered care. The current and future potential of mHealth is then examined for specific areas of allergology, including allergic rhinitis, aerobiology, allergen immunotherapy, asthma, dermatological diseases, food allergies, anaphylaxis, insect venom, and drug allergy. The impact of mobile technologies and associated big data sets are outlined. Facts, recommendations, and an action plan for future mHealth initiatives within EAACI are listed.

1. Becoming Wireless – The Frantic Evolution of Information and Communication Technologies

Today's world is connected wirelessly. This is reflected by the fact that the amount of mobile phone subscriptions has overtaken the number of people on the planet, a phenomenon being accompanied by a growing number of broadband connections for these phones, which creates a ubiquitous mobile infrastructure [1]. This chance has been seized by a multitude of companies, developers,

private entrepreneurs and start-ups, which has created an avalanche of mobile applications (Apps) with services mainly focused on entertainment, infotainment and the ease of daily life procedures. Interestingly, the development of healthcare related Apps and devices represents the fastest growing area within the information and communication technology (ICT) sector. This offers immense opportunities for global healthcare systems facing the challenge of improving patient care by making it more precise, efficient and cost-effective while improving accessibility especially for remote areas. To date, most of the evolution in the Mobile Health (mHealth) sector has been driven by private companies but central structures to ensure the quality of existing and new products have not yet been established. This urgent need has been recognized by the World Health Organization (WHO) [2], the European Union, national governments and a multitude of medical associations [3].

As millions of patients suffering from allergic diseases may benefit from mHealth innovations, the European Academy of Allergy and Clinical Immunology (EAACI) created a task force to assess the state of the art as well as the future potential of ICT in the field of allergy. The evaluation of 136 mobile applications in 2016 depicted a broad heterogeneity in terms of content and quality. As the mobile health environment is a very dynamic field, some of these may no longer exist or comply with regulatory requirements. Very few apps had been clinically validated and many were not based on guidelines or clinical evidence. Since then, various studies have evaluated the advantages, usability, efficiency and risks of mobile health technologies in allergic rhinitis [4-6], asthma [7-9], atopic dermatitis [10], food allergy [11, 12] and anaphylaxis [13].

Recognizing this scenario, the EAACI Task Force has created a position paper, summarizing general aspects such as legal regulations and evaluation criteria before evaluating the role of mHealth technologies in the respective allergic diseases. Finally, a roadmap for future actions of EAACI for the improvement of patient care through mHealth strategies will be depicted, considering possible limitations. A selected list of allergy-related Apps will be given in the appendices.

2. Endorsement of mHealth policy by WHO and American College of Allergy, Asthma and Immunology

2.1. "Be He@lthy, Be Mobile" - a WHO Initiative

The "Be He@Ithy, Be Mobile" (BHBM) initiative is a global partnership led by the World Health Organization (WHO) and the International Telecommunication Unit (ITU) [14]. It supports scale up of mHealth within national health systems to help combat diabetes, cancers, cardiovascular and chronic respiratory diseases. As mobile technologies and internet access are also widely spread in

countries with low average incomes, WHO recognized mobile health technologies as a valuable tool in providing healthcare to populations in remote areas or with limited access to health infrastructure. Training and self-empowerment become especially important in these cases. The handbook "mBreatheFreely" refers to the use of mobile technology to provide health information and support for people living with asthma and COPD. It provides guidance for governments and policymakers to develop, implement, and evaluate an mBreatheFreely program for the prevention and control of both diseases. The health messaging provided uses evidence-based behavior change techniques to help persons at risk of or affected by asthma and COPD to prevent and manage these conditions.

In addition to guidance for the implementation of concrete programs, WHO also initiated an mHealth Technical Evidence Review Group. Together with a panel of external experts, this group created a check-list of 16 items to standardize and improve the quality of mHealth evidence reporting [15].

2.2. Telemedicine in Allergy (Position Paper of the American College of Allergy, Asthma & Immunology)

The American College of Allergy, Asthma and Immunology (ACAAI) created a task force to evaluate the advantages and limitations of digital technologies within the broader scope of telemedicine [3]. In summary, ACAAI considers telemedicine a valuable method for health care delivery, especially to patients in rural or remote areas. The paper states that it may enhance patient-doctor collaborations and improve adherence as well as health outcomes. By facilitating access to specialists, it is valuable especially for allergic patients whose condition often requires prompt assessment, a need often hindered by long waiting lists for appointments with specialists. Although underlining the strength and positive potential of eHealth, the authors mention various challenges, such as standardized regulations, privacy, security, licensing, credentialing and reimbursement. This position paper gives a valuable general background for the evaluation of mHealth services in allergy care.

3. Regulatory Background

3.1. Legal Background for Centralized Quality Control and Risk Management

To access markets, minimize risks and gain relevance by providing the necessary level of trust, mHealth services must meet requirements of numerous legal domains. Unfortunately, many of these

lack international harmonization. For instance, medical liability and remote treatment are not covered by multi-national treaties or acts and differ across Europe. Thus, an international standard covering all legal aspects of m-Health in detail is not achievable. However, certain legal areas offer instruments for creating multi-national standards. For example, the "General Data Protection Regulation" [16] incentivizes the development of domain-specific Codes of Conduct by providing proof of compliance within the whole of Europe; an example addressing mHealth can be found online [17]. Any standardization initiative should use such tools extensively. To reflect the importance of full compliance in non-harmonized legal domains, those initiatives should additionally identify and implement equal international requirements as far as possible - supplemented by the obligation of a full legal compliance assessment for each territory in which a service is provided. In addition, technologies incorporating medical diagnosis and intervention should be registered as a medical device (Medical Device Directive 43/42/EWG) and obtain CE certification (CE1, CE2). Regulations and their interpretations are evolving and as such, any recommendations will need to be regularly updated.

3.2 Evaluation Criteria for mHealth Tools

Any medical intervention or tool, including mHealth Apps, may have potential risks and benefits [18] Several tools to assess Health App quality have been developed. The first methods covered mainly usability [19] while others focused on the development and life cycle of the App [20]. The Health Care Information and Management Systems Society published a guideline to evaluate the usability, but did not include any information quality criteria [21]. Newer tools have been developed, informed by systematic reviews of the literature and validated for internal consistency and interrater reliability, such as MARS [22], U-MARS (an end-user version)[23] and Enlight [24]. MARS and U-MARS evaluate engagement, functionality, aesthetics, information (including credibility) and subjective quality. Enlight also incorporates domains to assess therapeutic potential and tool generalizability. A recent systematic review identified relevant quality domains of user-facing e-Health programs such as usability, visual design, user engagement, content, behavior change/persuasive design, influence of social presence, therapeutic alliance, classification, credibility/accountability, and privacy/security, and highlights a high degree of agreement on these criteria around the globe [25]. Several institutions are beginning to offer mHealth accreditation.

4. Stakeholders' Role

4.1. The use of mHealth technologies from the patients' perspective

Patients may use mHealth for multiple reasons, which may not be aligned with the aims and objectives of clinicians [26] "Patient centred care should be personalised, pro-active and patient driven" [27]. Health care needs and wants have to be differentiated. App design should incorporate facultative use to enable tailoring [28]. To ensure patient centeredness and relevance, patients should be involved at every phase of the design, implementation and updating process [15]. For patient management, apps could be offered as part of information giving, monitoring and self-management to facilitate patient participation. Patients need to have confidence in the validity of any app used, thus the provenance should be explicit [29,30].

4.2. The use of mHealth technologies from the healthcare professionals' (HCP) perspective

HCPs are co-end users of mHealth, and only then, with patient assent. The mismatch of priorities between patients and clinicians need to be reconciled prior to developing any mHealth intervention to create a therapeutic partnership between patient, clinician and mHealth [15]. Secondary concerns of HCPs include incorporating data into electronic health care records and clinical responsibility for their use [18]. HCPs attitudes and systems (IT, organizational and incentivisation) will need significant reorientation to incorporate mHealth into routine medical care, which will require permissive cultural and organizational changes [31,32].

mHealth is an integral part of clinical care as an auxiliary feature aimed at improving quality of care, patient outcomes, and delivering efficiencies. The immediacy of app interaction and how this is provided (algorithmically or personally) needs to be addressed. Research is needed to understand the patterns of patient usage of Apps as well as the impact of mHealth technologies that require a prescription or formal physician oversight (digital therapeutics). Within the context of personalized precision medicine, mHealth Apps could facilitate the change of the model of care from clinician- to patient-centered care [33].

5. mHealth in Allergic Diseases

5.1. Allergic Rhinitis

Currently, the impact of mHealth on the diagnosis of rhinitis is small, with a limited number of mHealth tools for allergic rhinitis (AR) diagnosis published in peer-reviewed journals [34-36, 5]. Several others, from which published data is pending, are available on the market. Recent advances

in integrated biosensors, wireless communication and power harvesting techniques are spawning a new breed of point-of-care devices. However, AR is a very common disease and any diagnostic device connectable to a smartphone (e.g. peak nasal inspiratory flow meters, intranasal biosensors) will need to be very cheap to be affordable.

The monitoring of the control of allergic multimorbidities (rhinitis, conjunctivitis and asthma) has in contrast been approached by several apps. *Allergymonitor*, for example, allows the monitoring of symptoms and medication intake, which is then matched to local pollen concentrations [36, 37, 5]. The *MASK* (Mobile Airways Sentinel Network) *MASK-Air*, initially called *Allergy Diary* uses a visual analogue scale (VAS) for nose, eye and asthma symptoms, work impairment and a global assessment [38]. The data collected by the users of this app has led to new insights on work productivity, treatment patterns or phenotypes of allergic diseases.

Another promising aim for mHealth tools is improving our understanding of how patients adhere to medication adaptively. Lack of understanding of medication usage is common in all chronic diseases. Studies of patients using the *MASK* App show that users' behavior is often not in accord with guidelines, but patient frequently treat themselves as needed, which results in only less than 5% taking medication according to guideline recommendations [39]. These results prompted ARIA to develop a self-management strategy rather than targeting an increase of adherence. Nevertheless, internet-based telemonitoring improves the taking of intranasal corticosteroid (INCS) and improves disease knowledge among children and adolescents with seasonal allergic rhinoconjunctivitis [6]. Push notifications offer a promising strategy for enhancing engagement with smartphone-based health interventions in allergic rhinitis.

Mobile Apps also have the potential of discovering new allergic disease patterns through the acquisition of large data sets. For example, *MASK* unearthed novel patterns of allergic multimorbidity, which had not been demonstrated in a previous study (Mechanisms of the Development of Allergy MeDALL) [40]. When applying this new information, novel patterns (asthma, rhinitis and conjunctivitis) could then be confirmed on re-analysis of MeDALL data [41].

5.2. Pollen, Fungal Spores and Aerobiology

People affected by pollen allergy need accurate pollen information/forecasts to assist allergy diagnosis, allergen avoidance and symptom management, thus improving quality of life [42,43]. Monitoring and forecasting should not be limited to pollen or spore concentrations, but also include

other environmental information such as ozone levels, sulfides, nitrogen dioxide, particulate matter and others, as these agents, in addition to their non-specific effects, may enhance pollen allergenicity [44].

The assessment of pollen and spore levels may aid the identification of clinically relevant allergies to specific plants or fungi as well as guiding decisions concerning allergen immunotherapy [36].

Nowadays, allergic symptoms may be documented electronically in pollen diaries (e.g. [34, 5]). The advantages of such online diaries are comprised of continuous monitoring of allergic symptoms, enabling comparisons of different years/seasons and with different aerobiological particles, among them pollen and fungal spore concentrations. Users learn more about their symptomatic pattern, track down a possible pollen allergy (when compared with pollen concentrations) and thus may be willing to visit an allergist/medical doctor earlier asking for advice, diagnosis and treatment. In the future, the combination of tracking symptoms and evaluating the personal exposure (outdoor/indoor exposure) will play a role.

However, certain quality criteria (e.g. the inclusion of pollen data elaborated by institutions capable of monitoring/assessing and evaluating aerobiological data bearing the scientific and ethical responsibility) should be defined especially concerning accurate pollen and pollutant forecasts and their incorporation into mHealth for pollen allergy sufferers to maximise benefits [45].

5.3. Allergen Immunotherapy

mHealth technology, including telemonitoring, integrated care pathways (ICPs) and clinical decision support systems (CDSS) are suggested as potential tools to aid decision-making for AIT, as well as the identification of clinical responders to treatment [46-48]. If algorithms are based on evidence-based clinical recommendations for AIT such as outlined in the EAACI guidelines [49-51], this technology has the potential to optimize the precision for prescriptions [36], as well as efficacious and evidence-based products in AIT. When AIT is initiated, mHealth technology may in addition effectively increase patients' adherence [52], which is reported to be low in AIT [53, 54]. Patient support programs (PSPs) have suggested improving adherence by integrating and optimizing communication, educational, motivational and behavioral modification components [55, 56]. These could be implemented in mHealth technology, for example electronic reminder systems, e-communication channels, the use of "push"- messaging, gaming, including social networks with caregivers and peers

[55]. mHealth telemonitoring is a promising tool to monitor clinical benefits and side-effects of AIT including improvement of symptoms and quality of life or medication reduction. These technologies are already in use, e.g. as e-diaries in clinical trials of AIT aiming to collect clinical data in real-time for research and AIT product development [57]. In addition, real-life monitoring of large populations of patients receiving AIT in routine clinical practice both during treatment and after treatment cessation ('carry over'-effect) may become possible with mHealth technologies. Additionally such large data sets offer the potential of identifying unmet needs to be investigated in the future [47]. These may include prospective evaluation of adherence in a real-life population and long-term clinical effects after cessation of AIT (which is not feasible in randomized controlled trials for ethical reasons, costs and patients' willingness) or pharmacoeconomic evaluations. The latter is of great importance for payors and health-systems.

5.4. Asthma

mHealth provides not only tools to support patients with asthma in self-monitoring and decision-making, but also offers a variety of digital therapeutics to support disease management [58,59]. In fact, mHealth has the potential to enhance the quality of care, improve adherence to therapy and detect deterioration of symptoms by continuous monitoring and feedback to patients. A meta-analysis demonstrated improved asthma control with the use of mHealth, though the quality of Apps was substantially heterogeneous [28].

Many asthma Apps have been developed and are available for use [60], mainly by adults, but some also for school-age children and adolescents [7, 61-64]. Registering the use of a reliever inhaler has been used to monitor pediatric asthma control and to provide feedback through an electronic treatment plan [7].

The importance of users' feedback has been underlined by the outcome of a project including adolescent volunteers (13-18 years-old) who evaluated two asthma apps (AsthmaMD and Asthma Pulse) [62]. The suggested improvements included push-reminders (to take medication and to purchase refill), asthma-related games, fun-factors and a built-in flow meter [63]. Recording clinical and functional endpoints (i.e. symptoms, FEV1, PEF) on a daily basis, together with allergen and pollutant exposure, facilitates continuous asthma monitoring. Smartphone-based technologies for the assessment of objective parameters, such as lung function or lung sounds, have been developed and are currently being evaluated. Receiving mHealth data prospectively in addition to history taking may improve diagnostic precision. mHealth can support tailored asthma patient education, provide

reminders and improve self-management (e.g. trigger avoidance, use of rescue therapy and behavioral guidance during exacerbations). Existing randomized-controlled trials (RCT) have mainly focused on comparing the effect of Apps on asthma control to paper-based asthma management [64, 65].

However, there is lack of long-term RCTs of mHealth for the improvement of asthma control [66]. With regard to this, hopefully, results from the recently completed multicenter Horizon 2020 EUfunded project "My Air Coach", aimed at developing an innovative asthma monitoring system, will be able to answer current unmet needs in the field [9].

5.5. Dermatological Diseases

Mobile Health can play a role in the care for patients with dermatological allergic diseases, such as atopic dermatitis, contact dermatitis, chronic urticaria and cutaneous manifestations of drug hypersensitivity. Once the diagnosis has been confirmed, Apps can be useful for the monitoring of complaints and other symptoms, the support of patient self-management, the facilitation of professional-patient communication, telemedicine and peer support or research.

The severity and extent of disease can be measured over time for better self-control of the disease in form of a patient diary. Using validated questionnaires, which are available in Apps, a graphical display of scores over time is shown and patients or caregivers get insight into the course of the disease and the effect of the use of medication or topical therapy. There are several validated instruments for scoring severity of dermatological diseases [67, 68], and the Patient Oriented Score of Atopic Dermatitis (PO-Scorad) has been deployed for use in a mobile App [69] as well as the Atopic Dermatitis Activity Score and the Patient Oriented Eczema Measure of the University of Nottingham [70]. Other specific tools measure the impact of chronic skin diseases on sleep quality, using wearable sleep and/or itch tracker. Medication reminders or adherence apps remind patients to use their medication in time and might help to support action plans [71].

Apps including information about the disease, playful information for children, treatment, living with the disease, videos and patient stories can support self-management of patients. Patient portal Apps, which allow patients to view their medical file, send e-consultations and request e-repeat prescriptions may facilitate patient-doctor communications [72-74]. This may also be supported by Apps to share photos between the patient and HCPs as well as between doctors for teledermatology. Automated image recognition may deliver additional support for professionals.

mHealth can also gather data for research purposes and support communication within patient groups.

Skin test results can be assessed and recorded on the skin and in clearly positive or negative outcomes could also be evaluated by morphometric analysis, documented, followed-up and shared by apps. Computer- or mobile-based morphometric analysis is easier regarding the erythema as compared to the wheal associated with positive skin test reactions, because colour changes (erythema, blanching due to compression of vessels in the wheal, reflections by vesicles/blisters in the patch test) are easier to detect than the swelling of the wheal. Digital photodocumentation of skin prick test, patch test and intradermal test results could be collected and analysed by mHealth. The principal correlation between doctor-based and computer-based morphometric evaluations of positive skin test responses has been reported [75]; however, no program or algorithm has been proposed for practical use yet.

5.6. Food Allergy

In a recent study a total of 77 food allergy apps were analyzed [76]. While some of them exclusively provide information (24.6%), the majority (67.5%) includes various tools, such as food scanners (27.5%), food diaries (23.5%), and symptom trackers (21.5%). Only six Apps contained both food allergy education material and tools. Additional features included allergy-friendly restaurant locators and educational games for children. However, no App enabled the creation of a personalized Food Allergy Action Plan generated by a specialized HCP. The authors concluded, that most of the food allergy Apps examined offered an incomplete spectrum of information for patients [76]. In contrast to other fields of allergy, no studies have been performed in order to evaluate the benefit of food allergy related mHealth technologies.

Mobile health in food allergy may play a role for different stakeholders including patients and patient organizations, doctors and allergy organizations but also the food industry [77—83, 11, 12]. Within food allergy different levels of medical management can be approached by mHealth tools. At the level of diagnosis mobile health tools can support patients for the documentation of symptoms. With respect to the labeling of allergenic substances in food items, EU legislation has provided a list of food allergens, which are required to be labelled. Barcodes are already used for food labeling and support patients for the identification of appropriate products. Apps dedicated to the identification of declared allergens in food products [e.g. ShopWell®, ipiit® and others] are widely distributed, but lack validation and often do not declare their source of information. Other Apps support allergy

patients in the selection of appropriate products, based on their specific allergen profile [e.g. FoodMaestro App®]. In case of an accidental contamination during food processing, effective alert systems for patients, are desirable. Further, tools translating food names into images or other languages are useful for food allergic patients when travelling to countries where their native language is not spoken.

Other mobile health applications in food allergy support self-management for acute reactions. These tools are similar to those used for anaphylaxis in general and will be discussed below.

Taken together, mobile health could have a significant impact on the management of food allergy. However, clinical validation of high quality tools is necessary before their distribution in order to avoid over-diagnosis and the occurrence of avoidable reactions due to inaccurate information. Close collaboration between the different stakeholders and further research are urgently needed.

5.7 Anaphylaxis

Currently mHealth tools are primarily used in patients with anaphylaxis for educational and interventional purposes [84, 13]. Potential stakeholders are patients and patient organizations as well as doctors and allergy organizations. Educational materials can increase the knowledge among the above mentioned target groups but also other individuals like teachers, nurses, preschool personnel and family members or other persons which may need to act during an anaphylactic reaction. The recognition of key symptoms can be supported by offering visual examples (photos, videos). The performance of acute treatment measures, especially the application of an adrenalin auto-injector (AAI) can be enhanced through anaphylaxis action plans delivered via mHealth technologies. This support may be implemented through direct audio instructions or automated emergency calls. Novel alert systems to identify carriers of epinephrine auto-injectors and emergency departments in proximity are currently in development and require a further evaluation.

Automatic alerts signaling to the patient the expiration of his/her adrenalin auto-injector have already been successfully used [85,86]. As anaphylaxis is life threatening but rare, the development and implementation of mobile training tools is of high priority to facilitate repeated training and thus optimize the competence of patients, doctors and other persons involved. Large amount of materials are available [87, 88]. A recent clinical trial on the use of a smart-case for epinephrine autoinjectors showed improved satisfaction related to decreased anxiety among patients using the device [89]. Further, participants reported on improved adherence to carrying the injector and

better involvement in anaphylaxis management. Despite these promising results, future research needs to include prospective clinical trials assessing the improved clinical outcome of anaphylactic patients within the context of mobile health tools.

5.8. Venom Allergy

Although mHealth technologies may be very useful for prevention and management of venom allergy, the number of existing applications is limited. Apps could be used to graphically report the presence of different Hymenoptera species or non-common species in certain regions. Hikers or travelers may use them to make pictures of species, which will be automatically identified and reported. This can be used as a base for warning systems of potentially dangerous insects. Moreover, mHealth can serve to communicate with emergency departments or authorities in isolated regions or when no help is present in case of a potentially severe allergic reaction [13]. mHealth also might help to record and identify the culprit insect after a stinging event. Hence, all these possible applications could improve the identification of Hymenoptera venom-allergic patients and could contribute to the prevention of severe reactions. With regard to the recognition and management of acute reactions, please refer to the chapter on anaphylaxis (5.07).

Furthermore, mobile applications should be developed to monitor the treatment course of venom immunotherapy (VIT), including dosage, local or systemic adverse reactions and reminders of the subsequent appointment to receive the next dose. Studies are needed to assess whether mHealth may also improve adherence and make patients co-responsible for their own treatment, as well as increase the awareness of the importance and suitability of venom immunotherapy.

5.9. Drug Allergy

mHealth Apps for drug allergy have been developed mainly for educational purposes to help distinguish adverse drug reactions between those which are pharmacologically explicable and those due to immediate or delayed hypersensitivity [90].

Very few apps specifically dealing with drug allergy have been developed. Therefore, there is an urgent need for applications providing information in the following areas: different manifestations of drug hypersensitivity, drug interactions/cross-reactivity, common differential diagnoses, frequent elicitors of different types of drug hypersensitivity and a list of brand names indicating related generic drugs in different countries. It should also include suggested therapeutic alternatives when a drug or class of drug is implicated. Quality control in the development of Apps is especially relevant

in the field of drug allergy as unintended use of drugs the patient is allergic to is quite common and reactions can be potentially life threatening. Apps aimed at the distinction between hypersensitivity reactions and those caused by other mechanisms are not recommended for being used by patients, as this requires specialized professional assessment. Also the re-evaluation of previously recorded but possibly yet unconfirmed drug allergies, may be assisted by digital health technology.

5.10. Complementary and Alternative medicine

Apps on Complementary and Alternative Medicine (CAM) have been developed, promising allergy relief with practices such as Acupressure and Hypnotherapy, but also diagnosis (e.g. detection of food sensitivities with a compatible heart monitor via "Bulletproof Food Detective"). EAACI has expressed opposition to unconventional diagnostic tests and discourages their use [91, 92]. Products and methods of CAM are not free of adverse-effects [93]. A competent mHealth App should be in accordance with evidence-based medicine, thus the use CAM Apps is not indicated.

6. Research

In addition to the transfer of information between patient and HCP, mHealth technologies entail new opportunities for research, especially epidemiological studies. These will profit greatly from the integration of real-life patient experience with increased technical savvy.

Mobile health technology offers enormous possibilities for allergy research in several aspects: epidemiology, surveillance, health economics, public-health, clinical diagnosis, monitoring therapy.

- Epidemiology: Data collection through Apps allows extremely rapid collection of data from populations of allergic patients; this will tremendously increase the dimensions of epidemiological studies in all areas of medicine, including allergology;
- Surveillance: the use of electronic clinical diaries make the daily monitoring of symptoms of huge amounts of patients possible, allowing easy and cost-effective real-life studies on the use and efficacy of drug therapy and allergen immunotherapy; additional data on pollen and spore concentrations give the base for establishing individual exposure-symptom-thresholds.
- Health Economics: Apps dedicated to monitoring patients treated in real life conditions will allow rapid and valid collection of data for health economic studies aimed at measuring the economic impact of new and old diagnostic procedures and treatments;
- Public-Health: Allergy Apps offer the possibility of daily monitoring the population of patients, whose position in a given administrative area is identifiable with geo-localization tools; this

possibility will facilitate the development of public health programs aimed at managing pollen allergy and other diseases whose symptoms are triggered by environmental factors; this will open-up opportunities to treat pollen allergy at community level, thus improving the cost-benefit ratio of allergy care in the population;

- Clinical Diagnosis: mobile Health has great potential to improving allergy diagnosis in this new Era of Precision Medicine; for example: the use of electronic clinical diaries allows matching the data of the individual patient with the trajectories of environmental triggers registered by public agencies, identify the patterns of triggers relevant for the patient and implementing appropriate and personalized prevention strategies;
- Apps linked to diagnostic device: smart phones are becoming the conveyor of objective data acquired by all sorts of diagnostic devices and biosensors; Apps integrate these data with other information acquired or entered by the patient and allow a steady monitoring of the patient symptoms and parameters; research in this area will change the way of advancing diagnosis of allergic diseases.

7. Facts and Recommendations

The advantages and opportunities illustrated above in the management of the allergic patient is counterbalanced by a long list of barriers. EAACI takes these challenges seriously while planning activities in this novel area of medicine:

Patient-Doctor Relationship

Facts: mHealth technologies offer valuable possibilities of communication and consultation even outside regular office hours. Furthermore, delocalization of the patient's data could facilitate remote second consultations with allergy specialists.

Recommendation: Direct and close contact between HCPs and patients (blended care) is fundamental for good patient care and should never be totally replaced by digital technology.

Quality control (medical + technical)

Facts: Patients and HCPs will be increasingly encouraged to use Allergy Apps whose quality, safety, efficacy, reliability and appropriateness is not verified by any public health authority or scientific organization. It is also often difficult to evaluate the technical appropriateness of Apps and related devices connected to the smartphone.

Recommendation: the CE certification as a medical device should always be a pre-condition for the certification or distribution of an Allergy App. Still, a certification does not free the physician of the responsibility to monitor the use and data outputs of applications.

Legislation

Facts: European (and Non-European) regulations on mobile health technology are growing in number, relevance and heterogeneity (see section 3.01).

Recommendations: Apps certified or produced by EAACI must respect recent rules (EU - General Data Protection Regulation) established at European level and their future upgrade. Moreover, country-specific rules will have to be taken into account at local level.

Licensing

Facts: the use of mobile health and telemedicine in the management of the allergic patient allows delivery of remote care by doctors who may have no license or credentials to practice as a doctor in general or even an allergy specialist where the patient is living.

Recommendations: medical licensing systems need to be adapted to this new situation.

Privacy and confidentiality

Facts: privacy and protection of sensitive data is one of the most common weak points of allergy Apps available on the digital market.

Recommendations: EAACI will not recommend the use of allergy Apps that are not compliant with the current European and local legislation on this matter.

Data Overload

Facts: Although the easy and rapid collection of large data sets is a great advantage of mHealth technologies, the processing and evaluation of these data represents a significant challenge for HCPs.

Recommendations: App developers should pay attention to this fact and integrate solutions for manageable data sets including incorporation into EMR (electronic medical/health records). To provide continuous and safe care, further actions related to the interpretation of acquired data need to be planned carefully in advance.

Ethical prerequisites

Facts: The rapid development of mHealth technologies enables external persons, companies and institutions to access the private sphere of a multitude of users. This accessibility does not only facilitate data collection, but also interventions. Both action require consideration of ethical aspects.

Recommendations: Apart from legal aspects, EAACI emphasizes that any development and performance of Apps requires careful ethical consideration.

Reimbursement

Facts: In most countries, the time and expertise spent by doctors and specialists in assisting their patients through Apps or other telemedicine tools is not paid; this limits more rapid adoption of new technology. Health insurance companies/systems and public administration are slowly acknowledging this problem but reimbursement practices are in their infancy and rather sparse and episodic.

Recommendations: The use of validated mHealth tools should be reimbursed if used for improved care in the clinical practice of doctors.

Interference with disease management plans

Facts: improper use of Apps and other telemedicine tools may threaten the continuity of the relationship between the patient and his doctor, increasing the tendency to inappropriately self-care unguided by a proper disease self-management plan.

Recommendations: doctors should be aware of this risk and address it directly with their patients. Both parties should know the Apps and devices used by the patient. mHealth should be established as a form of blended care within any integrated care pathway.

Interoperability

Facts: The harmonization of different data management systems is a significant challenge for IT developers and HCPs.

Recommendations: The integration of mHealth data into electronic health records, e.g. at hospitals, outpatient clinics or within primary care is fundamentally important to ensure continuity of care.

Accessibility

Facts: A non-negligible proportion of the European population does not have access to a smartphone nor has sufficient health and digital literacy [94].

Recommendations: The experience acquired by WHO programs on mHealth in low- and middle-income countries may be useful to face this challenge also in Europe.

Accreditation and Training

Facts: There is no accreditation system for the use by doctors of mobile health technology, nor is this area part of the curriculum for doctors or specialists; the level of awareness and education of doctors in the use of mobile Health technology is extremely low.

Recommendations: The correct and careful use of mobile health technologies and telemedicine tools should become part of the curriculum in the training of health care professionals in order to ensure an adequate level of awareness.

Research

Facts: Mobile health technology offers enormous possibilities for research. Published studies on the use of mobile health in allergic diseases are still very limited.

Recommendations: Research on the use of mHealth in allergic diseases requires urgent funding and expansion in every area, such as epidemiology, surveillance, health economics, public health, clinical diagnosis, monitoring therapy.

8. Conclusions

Allergology, as any other area of medicine, will be deeply influenced by mobile Health technology. Allergists and their patients have a new way of communication, through the phone camera, -sound recording system, -motion sensors, texting and ultimately by using diagnostic devices and diagnostic algorithms incorporated within the mobile phone itself. The revolution that these possibilities are bringing in epidemiology, care, and research has already arrived. The role of doctors, and in particular allergists, will be progressively altered. To contribute to this trend, the EAACI Task Force for mHealth and Allergy has designed a two-years long action plan (2018-2020) that will be implemented under EAACI leadership [e-repository]. Accordingly, EAACI recognizes the advent of the m-Health Era in medicine and contributes to its development proactively.

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 Table 1 Selected overview of allergy-related apps and their most important features (apps are categorized and sorted alphabetically)

	App (Name)	Purpose	Medical Information	Technical Characteristics	Additional Information
	Allergy- monitor	 Improvement of allergy diagnosis (matching SMS and pollen) Enhancement of patient-doctor communication Improvement of adherence to therapy and AIT Short-term prediction of patients' symptoms 	 Several Symptom and Medication Scores (RTSS, AdSS, RMS, ACS, dCSMS, VAS) Asthma score QoL score Matching SMS and pollen trajectories (AV) 	 Patient report (eye, nose, asthma symptoms, medication intake, side effects) Customized over-the-counter and prescribed medication list for all countries Patient-doctor communication via SMS and email Patient alerts for better adherence 	Number of Downloads: 5K+ Languages: 11 Costs: App-Free / Backend-Charged Website: http://www.tpsproduction.com/
Rhinitis	AllyScience	 Monitoring of allergic rhinoconjunctivitis and asthma symptoms Pollen information Community symptom reports 	 Monitoring of allergic rhinitis symptoms Pollen and symptom forecasts Geolocation Explicit data collection for study purposes 	 Users enter their 3 most attended locations Patient report (eye, nose, asthma symptoms) Community reports from other users Pollen information and forecast 	Number of Downloads: 1K+ Languages: 5 (only available in Switzerland) Costs: Free Website: https://allyscience.ch/
Allergic Rhinitis	MACVIA- ARIA MASK-Air (previously: Allergy Diary)	Assessment of symptoms and medication use in patients with allergic rhinitis	 VAS for allergic rhinitis, conjuntivitis, asthma and work productivity Recording of medication use WPAI-AS (Allergy-specific Work Productivity and Activity Impairment Questionnaire) Results on adherence to treatment, medications Clinical Decision Support System EQ5-D 	Customized over-the-counter and prescribed medication list for all countries	Number of Downloads: 15K+ Languages: 16 Costs: Free Website: http://www.allergydiary.com/all ergy-diary-app/
	Pollen App	 Allergen avoidance by the means of pollen information and forecasts Symptom information 	 Documentation of allergic symptoms and medication use Allergy risk assessment Therapy recommendations 	 Calculation of hourly allergy risk based on pollen forecasts Personal pollen information Symptom diary 	Number of Downloads: 100K+ Languages: 6 (with PASYFO: 9) Costs: Free

	Asthma Action Hero	Clinical Diary and Support for Asthma Patients	 Daily interactive plan including a detailed overview of how to identify moderate to severe symptoms of an asthma attack Clinical Diary of asthma symptoms 	 Plant encyclopedia Pollen alarm (Countdown for the season, push notifications for selected levels and aeroallergens) Symptom and medication log can be shared with the doctor Reminder system for medication intake 	Website: https://www.polleninfo.org/DE/ en/free-pollen-app.html Number of Downloads: 100+ Languages: 1 (English) Costs: Free Website: http://www.asthmaactionhero.c
Asthma	Asthma MD	Assessment of symptoms and medication use in patients suffering from asthma	 List of prescribed medication Clinical Diary of asthma symptoms Registry of Peak Flow Meter Measurements Correlation of asthma symptoms with environmental triggers 	Patient report can be shared with the doctor	om/ Number of Downloads: 10K+ Languages: 1 (English) Costs: Free
	Inspirer- mundi	Assess and promote: 1) adherence to preventive inhalers and 2) asthma control	 Therapeutic plan with personalised notifications; verified assessment of treatment adherence; monitoring of asthma symptoms and burden; asthma and rhinitis control (CARAT - Control of Allergic Rhinitis and Asthma Test) 	 detection of inhaler use (6 inhalers with dose counter) by processing of video images captured with the smartphone camera Three components: Monitoring (disease status and inhaler adherence), gamification (player starts as a "warrior" and evolves to become an "inspirer") and peer support Graphical display of monitored data Transmission of data to physician messaging system privacy and security features 	Number of Downloads: 100+ Languages: 2 (Portuguese, Spanish) Costs: Free Website: https://www.facebook.com/Proj eto-Inspirers- 218849795347948/

	My Air Coach	Support for Asthma Patients	 Sensors monitor and store several physiological, behavioural and environmental factors that will be crossed with asthma data. Creation of tailored asthma plans. 		Available only for Android Languages: 1 (English) Costs: Free Website: http://www.myaircoach.eu/
	My Eczema Tracker	For patients with atopic dermatitis	Eczema tracker based on the validated Patient oriented Eczema measure	Total symtpom score is calculated and scores are graphically displayed over time	Number of Downloads: 100+ Languages: 1 (English) Costs: Free Website: https://nationaleczema.org/wha t-is-measured-is-managed/
Diseases	PO-SCORAD	Self assessment to better monitor and better treat for patients with atopic eczema	Eczema severity score	Severity including itch can be recorded, developments will be shown in diagnams, patient can send a screenshot to the doctor, add photos	Number of Downloads: 5K+ Languages: 22 Costs: Free Website: https://www.poscorad.com/#/
Dermatological Diseases	Skin Safe	For patients with allergic contact dermatitis: lists products with contact allergens	Information about allergens in skin care products	 Creates an individualised allergen list through health care provider according to patch test results. Shows cosmetic products safe for the allergic patient After scanning a product: report on contained allergens relevant for the patient Information on hypoallergenic products, personalized list of favorite products helps to buy products directly from retailers 	Number of Downloads: 1K+ Languages: 1 (English) Costs: Free Website: https://www.skinsafeproducts.c om/apps Developed with the MayoClinic Lots of products, mainly on the American market

	Urticaria app	For patients with urticaria	Information and monitoring of urticaria	 Gives information on symptoms and dagnosis of urticaria, calculates the score for the urticaria control test, the 7-day urticaria activity score and the angeioedema activity score and creates a diary over time. 	Number of Downloads: 1K+ Languages: 2 (English, German) Costs: Free Website: https://play.google.com/store/a
				Developed by the Global Allergy and Asthma patient platform, which is sponsored by pharmaceutical companies including Novartis, several similar Apps by Novartis	pps/details?id=at.alysis.urticaria &hl=de
	Allergy Journal	Nutrition and Symptoms Diary	 Individual documentation of ingested foods and symptoms facilitate the interpretation of delayed reactions. Alert for offending foods. 	 Detailed reports to be sent by e-mail and exported on request as PDF or Excel Data storage on phone or linked dropbox 	Number of Downloads: ? Languages: 1 (English) Costs: iOS: 0,99€; Android: 1,09€ Website: https://itunes.apple.com/de/ap p/allergy-
Food Allergy & Anaphylaxis	Anaphylaxis	- Smartcase for epinephrine autoinjectors that connects wirelessly to a mobile application or works autonomously - Expiration date monitoring - Find Device Near me Map (pharmacies and hospitals) - parental control information for the	 Allows the user to know at any time the condition of his/her epinephrine. activates action plan in the event of anaphylaxis, giving clear instructions on how to administer the drug and notifying your pre-defined contacts of the incident, giving your exact location. Warns the user when too far away from the autoinjector Training for correctdrug 	 Report about the temperature and conditions of the medication (adrenaline) Controls transparency of Adrenaline GPS localization of the patient suffering an anaphylactic reaction and warns the preferred contacts that the patient is suffering an anaphylactic reaction Shows pharmacies and hospitals close to the user 	journal/id455223808?mt=8 Number of Downloads: < 5K Languages: 2 Countries: 27 Costs: Free Website: http://www.adanmi.com/anapp hylaxis/

e-symptoms	Electronical symptom diary	 Personal profile To observe, quantify and document symptoms, complaints and lifestyle habits Advice section Information material from the aha! Allergy Center Switzerland 	Symptom diary Informative section	Number of Downloads: 5K+ Languages: 4 Costs: free Website: https://www.aha.ch/
FoodMaestro APP®	Shopping advisor for patients with food allergy	 Multiple profiles for different family members Individual recommendation on healthy non-offending foods Shopping recommendations from a doctor or dietician can be saved Individual shopping lists 	 Barcode scanner with different nutrition preferences and allergy pre-settings Recommendations of products which suit the personal profile 	Number of Downloads: 46K Languages: 1 (English) Costs: free Website: https://www.foodmaestro.me/h ome.html
ipiit®	Shopping advisor for patients with food allergy	 Support for buying healthy groceries and managing food allergies. Information on the background of different recommendations Offering alternative products 	 Barcode scanner with preference selection (Gluten-free, Lactose-free and non-GMO). Compares products and ingredients - calories, fat, sodium, carbs, fiber, sugar, protein and iron. Community ratings 	Number of Downloads: 10K+ Languages: 1 (English) Costs: free Website: https://www.ipiit.com/
Lifesaver	This app allows the patients affected by anaphylaxis to become familiar with their illness and the handling of its auto-injector of adrenaline. It also incorporates tools that facilitate the patient's awareness to their environment on allergic reactions more serious.	It trains about the use of the adrenaline autoinjector Alerts about the date of expiration of the adrenaline autoinjector	Training on how to administer the drug	Number of Downloads: 100+ Languages: 18 Costs: Free Website: https://www.lifesaverapp.com/

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gy & Anaphylaxis	Rescufy	Connecting the patient and caretakers to medical help when anaphylaxis strikes.	 Connecting epinephrine injector to mobile phone including an alert if the user is too far away from the device Communicates vitally important allergy, medical and insurance information to Emergency Responders and emergency contacts 	 Captures information on allergies, any treatment previously administered, other medical conditions and insurance details Information on prescribed emergency medication Sends geolocalisation to key contacts and emergency responders automatic dialling of emergency number (911) when activated Patient alerts for better adherence 	Number of Downloads: 100+ Languages: 2 Costs: Free
Food Allergy	ShopWell®	Shopping advisor for patients with food allergy	 Personal profile Alerts about offending foods and provides recommendations on new/alternative shopping options available in the same store 	 Bar code scanner to find the ingredients in about 400,000 foods. Personalized food recommendations Location awareness to find products in your local grocery store S Health integration for personalized nutrition labels Buy recommended foods right from the app with our easy mobile shopping feature 	Number of Downloads: 100K+ Languages: 1 (English) Costs: free Website: https://www.innit.com/shopwell /