

University of Groningen

EUFOREA treatment algorithm for allergic rhinitis

Hellings, P. W.; Scadding, G.; Bachert, C.; Bjermer, L.; Canonica, G. W.; Cardell, L. O.; Carney, A. S.; Constantinidis, J.; Deneyer, L.; Diamant, Z.

Published in:
Rhinology

IMPORTANT NOTE: You are advised to consult the publisher's version (publisher's PDF) if you wish to cite from it. Please check the document version below.

Document Version
Publisher's PDF, also known as Version of record

Publication date:
2020

[Link to publication in University of Groningen/UMCG research database](#)

Citation for published version (APA):

Hellings, P. W., Scadding, G., Bachert, C., Bjermer, L., Canonica, G. W., Cardell, L. O., Carney, A. S., Constantinidis, J., Deneyer, L., Diamant, Z., Durham, S., Gevaert, P., Harvey, R., Hopkins, C., Kjeldsen, A., Klimek, L., Lund, V. J., Price, D., Rimmer, J., ... Fokkens, W. J. (2020). EUFOREA treatment algorithm for allergic rhinitis. *Rhinology*, 58(6), 618-622. <https://www.rhinologyjournal.com/Abstract.php?id=2664>

Copyright

Other than for strictly personal use, it is not permitted to download or to forward/distribute the text or part of it without the consent of the author(s) and/or copyright holder(s), unless the work is under an open content license (like Creative Commons).

The publication may also be distributed here under the terms of Article 25fa of the Dutch Copyright Act, indicated by the "Taverne" license. More information can be found on the University of Groningen website: <https://www.rug.nl/library/open-access/self-archiving-pure/taverne-amendment>.

Take-down policy

If you believe that this document breaches copyright please contact us providing details, and we will remove access to the work immediately and investigate your claim.

Downloaded from the University of Groningen/UMCG research database (Pure): <http://www.rug.nl/research/portal>. For technical reasons the number of authors shown on this cover page is limited to 10 maximum.

EUFOREA treatment algorithm for allergic rhinitis*

P.W. Hellings¹⁻⁴, G. Scadding⁵, C. Bachert⁷⁻⁹, L. Bjermer¹⁰, G.W. Canonica¹¹, L.O. Cardell¹², A.S. Carney¹³, J. Constantinidis¹⁴, L. Deneyer¹⁵, Z. Diamant¹⁶⁻¹⁸, S. Durham^{19,20}, P. Gevaert⁷, R. Harvey^{21,22}, C. Hopkins²³, A. Kjeldsen^{24,25}, L. Klimek^{26,27}, V.J. Lund²⁸, D. Price²⁹⁻³¹, J. Rimmer³², D. Ryan³³, G. Roberts³⁴⁻³⁶, P. Sahlstrand-Johnson³⁷, S. Salmi³⁸, M. Samji³⁹, Guy Scadding⁴⁰, P. Smith⁴¹, A. Steinsvik⁴², M. Wagenmann⁴³, S. Seys¹⁴, U. Wahn⁴⁴, W.J. Fokkens⁴

Rhinology 58: 6, 618 - 622, 2020

<https://doi.org/10.4193/Rhin20.246>

***Received for publication:**

May 19, 2020

Accepted: July 20, 2020

¹ KU Leuven Department of Microbiology, Immunology and Transplantation, Laboratory of Allergy and Clinical Immunology Research Group, Leuven, Belgium; ² University Hospitals Leuven, Department of Otorhinolaryngology, Leuven, Belgium; ³ University Hospital Ghent, Department of Otorhinolaryngology, Laboratory of Upper Airways Research, Ghent, Belgium; ⁴ Academic Medical Center, University of Amsterdam, Department of Otorhinolaryngology, Amsterdam, The Netherlands; ⁵ RNENT Hospital, Huntley Street, London, UK; ⁷ Upper Airways Research Laboratory, Dept of Otorhinolaryngology, Ghent University Hospital, Ghent, Belgium; ⁸ Division of ENT diseases, CLINTEC, Karolinska Institute, University of Stockholm, Sweden; ⁹ Sun Yat-sen University, International Airway Research Center, First Affiliated Hospital, Guangzhou, China; ¹⁰ Dept of Respiratory Medicine and Allergology, Skane University Hospital, Lund, Sweden; ¹¹ Personalized Medicine Asthma and Allergy Clinic, Humanitas University and Research Hospital, Milan, Italy, and SANI-Severe Asthma Network Italy; ¹² Division of Ear, Nose and Throat Diseases, Department of Clinical Sciences, Intervention and Technology, Karolinska Institutet, Stockholm, Sweden; ¹³ Ear, Nose, and Throat (ENT) Department, Flinders University, Bedford Park, South Australia, Australia; ¹⁴ 1st Department of ORL, Head and Neck Surgery, Aristotle University, AHEPA Hospital, Thessaloniki, Greece; ¹⁵ European Forum for Research and Education in Allergy and Airway Diseases (EUFOREA), Brussels, Belgium; ¹⁶ Dept of Respiratory Medicine & Allergology, Institute for Clinical Science, Skane University Hospital, Lund University, Lund, Sweden; ¹⁷ Department of Respiratory Medicine, First Faculty of Medicine, Charles University and Thomayer Hospital, Prague, Czech Republic; ¹⁸ Dept Clin Pharm and Pharmacol, Univ Groningen, Univ Med Ctr Groningen, Groningen, Netherlands; ¹⁹ Immunomodulation and Tolerance Group, Allergy and Clinical Immunology, Inflammation, Repair and Development, National Heart and Lung Institute, Imperial College London; ²⁰ MRC and Asthma UK Centre in Allergic Mechanisms of Asthma, London, United Kingdom; ²¹ Rhinology and Skull Base, Applied medical research center, University of New South Wales, Sydney, Australia; ²² Faculty of medicine and health sciences, Macquarie University, Sydney, Australia; ²³ Ear, Nose and Throat Department, Guys and St. Thomas Hospital, London, United Kingdom; ²⁴ Department of Otorhinolaryngology Head and Neck surgery, Odense University Hospital, Denmark; ²⁵ University of Southern Denmark, Odense, Denmark; ²⁶ Center for Rhinology and Allergology, Wiesbaden, Germany; ²⁷ Mainz University Allergy Center, Mainz, Germany; ²⁸ Royal National Throat, Nose and Ear Hospital, UCLH, London, UK; ²⁹ Optimum Patient Care, Cambridge, UK; ³⁰ Observational and Pragmatic Research Institute, Singapore; ³¹ Academic Primary Care, University of Aberdeen, Aberdeen, UK; ³² Monash Health, Monash University, Melbourne, Australia; ³³ Usher institute, University of Edinburgh, Edinburgh, UK; ³⁴ The David Hide Asthma and Allergy Research Centre, St Mary's Hospital, Newport Isle of Wight, United Kingdom; ³⁵ NIHR Biomedical Research Centre, University Hospital Southampton NHS Foundation Trust, Southampton, United Kingdom; ³⁶ University of Southampton, Southampton, United Kingdom; ³⁷ Department of Oto-Rhino-Laryngology, Head and Neck Surgery, Skåne University Hospital, Malmö, Sweden; ³⁸ Helsinki University Hospital, Helsinki, Finland; ³⁹ Imperial College London, London, UK; ⁴⁰ Royal Brompton and Harefield NHS Trust, London, UK; ⁴¹ Clinical Medicine, Griffith University, Southport, QLD, Australia; ⁴² Department of Otorhinolaryngology, Oslo University Hospital, Rikshospitalet, Oslo, Norway; ⁴³ Department of Otorhinolaryngology, Universitätsklinikum Düsseldorf, Dusseldorf, Germany; ⁴⁴ Klinik für Pädiatrie m.S. Pneumologie und Immunologie, Charite, Berlin, Germany.

To the Editor:

Allergic rhinitis (AR) is the most common chronic inflammatory disease, affecting an estimated 100 million Europeans⁽¹⁾. Despite a substantial burden on individuals, society and health economies⁽²⁾, AR remains under-diagnosed, under-estimated (in terms of severity), and under-treated⁽³⁾. Although effective

and safe treatments exist, patients wait too long to seek medical advice, often preferring to self-manage at drug stores and at the pharmacy⁽⁴⁾. Other barriers to access of appropriate and effective AR treatment exist at patient, pharmacist and physician levels, including inability to recognize AR and diagnose it, inappropriate AR medication prescription/use⁽⁵⁾, poor concordance

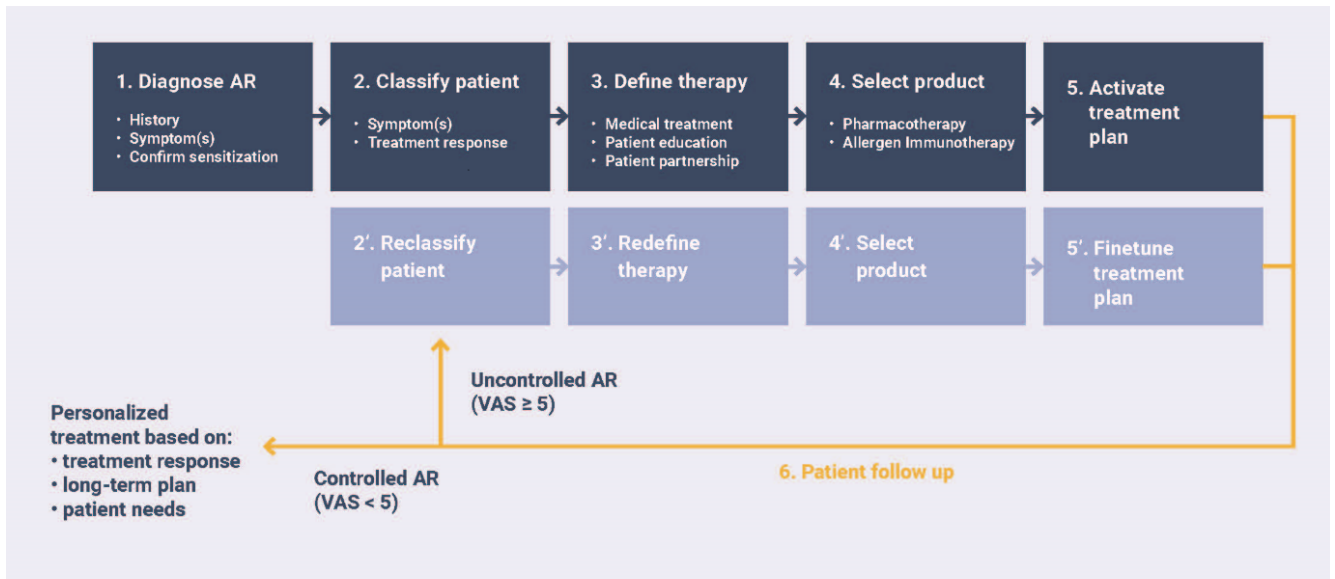


Figure 1. EUFOREA allergic rhinitis pocket guide implementation pathway. AR: allergic rhinitis; European Forum for Research & Education in Allergy & Airway Diseases; VAS: visual analogue scale.

with AR treatment regimens and/or lack of awareness of new medications. There has, therefore, been a shift towards a more patient-centred approach to AR management, with a focus on personalized, predictive, preventative and participatory strategies^(6,7). The visual analogue scale (VAS) was introduced as the common language of AR control, improving patient-healthcare provider communication, and informing disease control status and treatment recommendations⁽⁸⁾. However, guidelines based solely on VAS may not reflect the needs of physicians and patients in real-life, since VAS are not routinely used in everyday practice and may not capture the profiles of all presenting patients.

The European Forum for Research & Education in Allergy & Airway Diseases (EUFOREA) in collaboration with global key opinion leaders in the field of chronic inflammatory airway disease, has developed an AR pocket guide with a treatment algorithm to expedite access to AR treatment and facilitate coordinated care. The algorithm is designed for real-life use. Its aim is simple: to improve AR knowledge and streamline the transition of patients between self-, pharmacy-, GP- and specialist-care, allowing more coordinated care. The guide is practical and easy-to-use in everyday clinical practice for any care provider. It is concise and patient-centred, capturing every patient that attends the outpatient clinic of any care provider. This guide provides a 'what to do' checklist when assessing AR patients, including a list of symptoms suggestive of AR, questions on suspected asthma, and instructions on how to use the AR VAS. The AR pocket guide is implemented in 5 easy steps: (i) diagnose AR, (ii) classify patients, (iii) define therapy, (iv) select product, and (v) activate treatment plan (Figure 1). Diagnosis involves taking a comprehen-

sive history and investigating signs and symptoms, confirmed (if necessary) by identification of sensitizing allergen(s) linked to symptoms⁽⁹⁾. The patient is classified according to disease control and response to treatment using the AR VAS (retrospectively, if VAS is not already routinely monitored). Approach to treatment is defined, including discussion of potential benefits of allergen avoidance (whenever possible) and other provoking triggers, saline nasal sprays/douching and available treatment options. Patient education is central at all stages (e.g. disease information, awareness of symptoms, importance of adherence and correct use of intranasal sprays). Patient participation in the decision-making process and in goal-setting is encouraged, and therapy matched to these goals and to patient preference. AR treatment is selected depending on type and history of patient, disease control (assessed by VAS) and point of care (i.e. pharmacy, GP or specialist) (Figure 2).

Treatment Step 1: Patients with suspected AR presenting to any care provider. These patients should be treated with an intranasal corticosteroid (INS), non-sedating oral anti-histamine (OAH) or intranasal anti-histamine (INAH). Physicians' clinical experience and patient symptoms, preferences and expectations, provoking triggers and co-morbidities should be taken into account for optimal outcomes.

Treatment Step 2: Patients who have tried and failed (i.e. VAS score $\geq 5/10$ cm) Step 1 treatment at the pharmacy or previously at physician level. AR diagnosis should be confirmed, medication adherence checked, and co-morbidities evaluated. Treatment should be stepped up to fixed dose INS/INAH. Add-on therapy to INS is not recommended.

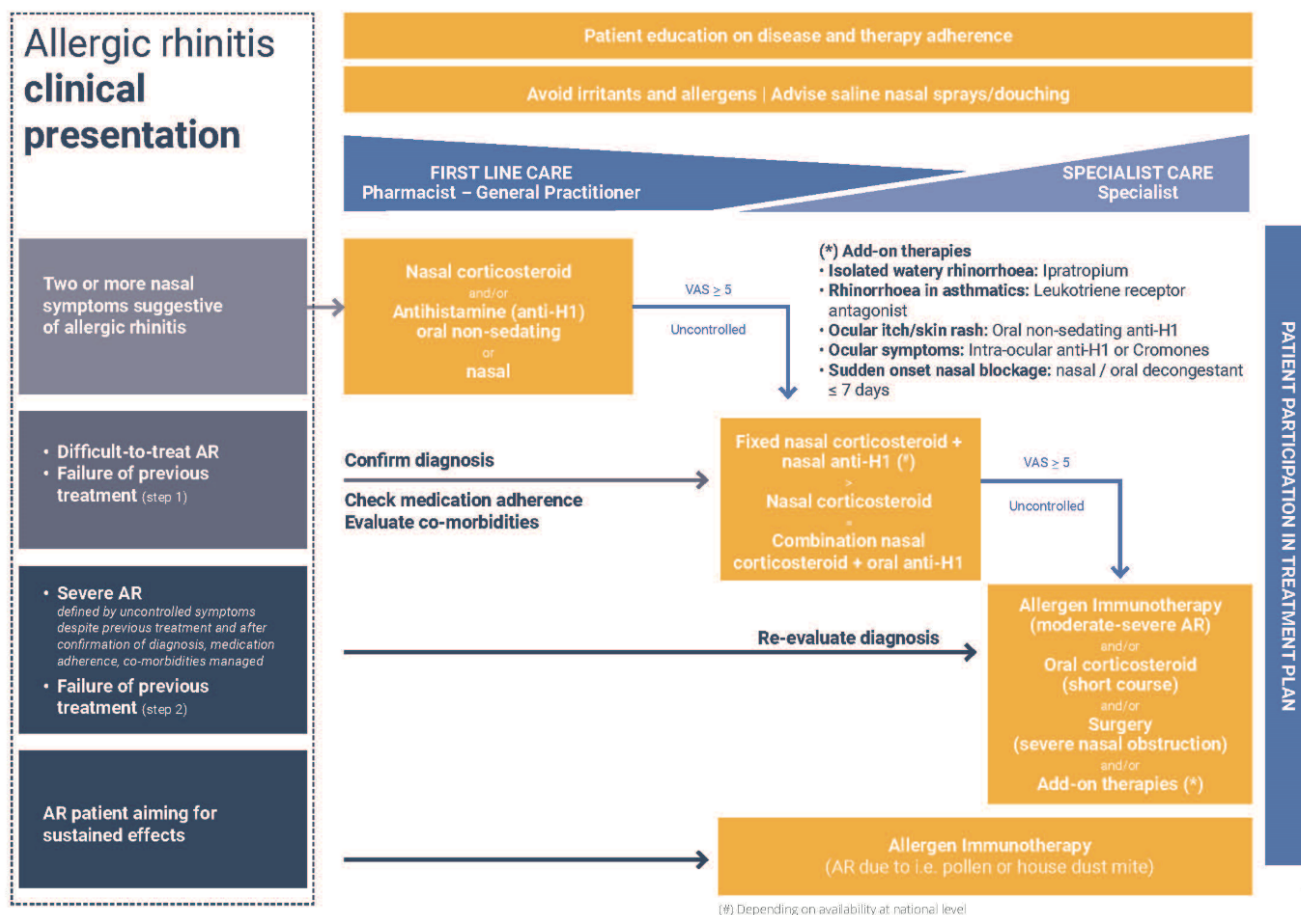


Figure 2. EUFOREA allergic rhinitis pocket guide treatment algorithm. AR: allergic rhinitis; VAS: visual analogue scale.

Treatment Step 3: Patients who have tried and failed Step 2 treatment (VAS remains $\geq 5/10$ cm) or those who present with severe symptoms. AR diagnosis should be (re-)evaluated, and symptom-directed add-on therapies to fixed dose INS/INAH considered (e.g. ipratropium, leukotriene receptor antagonist; non-sedating OAH, ocular anti-histamine/cromone and nasal/oral decongestant (≤ 7 days)) (Figure 2). Other Step 3 treatment options to consider include allergen-specific immunotherapy (AIT), short course OCS and surgery (for those with severe pharmacological therapy-resistant nasal obstruction), at physicians' discretion, considering availability, and risk/benefit ratio. AIT (subcutaneous or sublingual) is recommended for those patients looking for a sustained reduction of their rhinitis symptoms (if available and if appropriate to the patient's sensitization pattern, and their preference, lifestyle, adherence history and comorbidities (e.g. asthma)). AIT may be considered for patients with uncontrolled moderate-to-severe AR (+/- conjunctivitis) linked to exposure to allergens, with a confirmed IgE sensitization to these allergens and with inadequate control of symptoms despite pharmacotherapy and allergen avoidance measures and/or unacceptable adverse effects of medication. The next step involves the design and activation of a persona-

lized treatment plan by prescribing medication and explaining the expected response and treatment duration with the patient (Figure 1). It is essential that physicians explain the criteria for a return clinic visit (e.g. sustained VAS score $\geq 5/10$, adverse event). Use of digital technology to support adherence and to evaluate disease control may be suggested. A patient review (actual or digital) should be scheduled. If AR remains uncontrolled (i.e. sustained VAS score $\geq 5/10$ cm) despite completion of the implementation pathway (Figure 1), then AR needs to be re-classified, therapy re-defined, product re-selected and the treatment plan fine-tuned. The aim is to devise a treatment plan (with patient collaboration) which provides AR control, a treatment response acceptable to the patient, suitable for long-term implementation and in compliance with patient needs. The EUFOREA AR pocket guide with novel treatment algorithm designed for use in real-life, is concise, simple to use, suitable for all stakeholders including pharmacists, primary care physicians, ENT doctors, pulmonologists, allergists and paediatricians, and provides evidence-based and expert-endorsed AR management recommendations. This practical guide has the potential to ensure timely access to AR treatment, taking us one step closer to precision medicine, delivering the right treatment to the right

patient at the right time.

The full pocket guide is available on the EUFOREA website (<https://www.euforea.eu/>).

Conflict of interest

PWH: lecture fees and/or participation at expert board meetings of ALK, Stallergenes, Mylan, Novartis and Sanofi; GS: chaired the BSACI AR guidelines, has given paid lectures for and an education programme for EUFOREA. She also chairs the EAACI Ethics Committee, the Independent data monitoring committee for an ALK allergen immunotherapy trial and the Rhinology and Laryngology Research Fund and has given lectures for and/or advised ALK, Bayer GSK, Mylan, Stallergenes; CB: ALK, Stallergenes, Mylan; GWC: has received research grants, as well as lecture or advisory board fees from, A. Menarini, Alk-Abello, Allergy Therapeutics, Anallergo, AstraZeneca, Medimmune, Boehringer Ingelheim, Chiesi Farmaceutici, Circassia, Danone, Faes, Genentech, Guidotti-Malesci, GlaxoSmithKline, Hal Allergy, Merck, Merck Sharp & Dome, Mundipharma, Novartis, Orion, Sanofi-Aventis, Sanofi, Genzyme/Regeneron, Stallergenes, UCB Pharma, Uriach Pharma, Teva, Thermo Fisher, and Valeas; ZD: Apart from academic affiliations, ZD acts as Executive and Scientific Medical Director at a phase I/II pharmacological unit (QPS-NL), which performs clinical studies for pharmaceutical companies. In the past 3 years, ZD received honoraria, consultancy and speaker fees from Acucort, Astrazeneca, ALK, Aquilon, Boehringer Ingelheim, CSL, HAL Allergy, MSD, Sanofi-Genzyme; PG: has received lecture fees and/or participation at expert board meetings of Ablynx, ALK, Argenx, ALK, Astra-Zeneca, Genentech, HAL-Allergy, Novartis, Roche, Regeneron, Sanofi, and Stallergenes-Greer; RH: consultant with Medtronic, Olympus, Novartis and NeilMed pharmaceuticals. He has also been on the speakers' bureau for Glaxo-Smith-Kline, Meda Pharmaceutical, Seqiris and AstraZeneca. Research funding from Neilmed and Glaxo-Smith-Kline. CH: Advisory Board for Sanofi-Genzyme, Astra Zeneca, Olympus and Smith and Nephew; LK: has received research grants from Allergy Therapeutics/Bencard, Great Britain/Germany; ALK-Abelló, Denmark; Allergopharma, Germany; ASIT Biotech, Belgium; AstraZeneca, Sweden, Biomay, Austria, Boehringer Ingelheim, Germany, Circassia, USA; Stallergenes, France; Cytos, Switzerland; Curalogic, Denmark; HAL, Netherlands; Hartington, Spain; Lofarma, Italy; MEDA/Mylan, Sweden/USA; Novartis, Switzerland, Leti, Spain; ROXALL, Germany; GlaxoSmithKline (GSK), Great Britain; Sanofi, France and/or has served on the speaker's bureau or was consulting for the above mentioned pharmaceutical companies. LK is the current president of AeDA (German Society of Applied Allergology), a NAS of EAACI and Chair of the EAACI ENT section. VJL: has receive lecture and advisory board fees from Abbott, GSK, Johnson & Johnson, Novartis and Sanofi; DP: declares board membership with Aerocrine, Amgen, AstraZeneca, Boehringer Ingelheim, Chiesi, Mylan, Mundipharma, Napp Phar-

maceuticals, Novartis and Teva; consultancy agreements with Almirall, Amgen, AstraZeneca, Boehringer Ingelheim, Chiesi, GlaxoSmithKline, Mylan, Mundipharma, Napp Pharmaceuticals, Novartis, Pfizer, Teva and Theravance; grants and unrestricted funding for investigator-initiated studies (conducted through Observational and Pragmatic Research Institute Pte Ltd) from Aerocrine, AKL Research and Development Ltd, AstraZeneca, Boehringer Ingelheim, British Lung Foundation, Chiesi, Mylan, Mundipharma, Napp Pharmaceuticals, Novartis, Pfizer, Respiratory Effectiveness Group, Teva, Theravance, UK National Health Service and Zentiva; payment for lectures/speaking engagements from Almirall, AstraZeneca, Boehringer Ingelheim, Chiesi, Cipla, GlaxoSmithKline, Kyorin, Mylan, Merck, Mundipharma, Novartis, Pfizer, Skyepharma and Teva; payment for manuscript preparation from Mundipharma and Teva; payment for the development of educational materials from Mundipharma and Novartis; payment for travel/accommodation/meeting expenses from Aerocrine, AstraZeneca, Boehringer Ingelheim, Mundipharma, Napp Pharmaceuticals, Novartis and Teva; funding for patient enrolment or completion of research from Chiesi, Novartis, Teva and Zentiva; stock/stock options from AKL Research and Development Ltd which produces phytopharmaceuticals; owns 74% of the social enterprise Optimum Patient Care Ltd (Australia and UK) and 74% of Observational and Pragmatic Research Institute Pte Ltd (Singapore); and is a peer reviewer for grant committees of the Efficacy and Mechanism Evaluation programme and Health Technology Assessment; DR: Has received grants or payments from AZ,GSK,BI,Novartis, Regenron, Chiesi and Mylan; Ssa: has acted as paid consultant for ERT, Novartis, Sanofi Pharma, and Roche Products; MHS received research grants via Imperial College London from ASIT biotech, Regeneron, Merck, Allergopharma and UCB, and received lecture fee from ALK-Abelló, Allergopharma, Alletgy Therapeutic; MW: Fees for lectures or advisory boards: ALK-Abelló, Allergopharma, AstraZeneca, Bencard, Genzyme, HAL Allergie, LETI Pharma, MEDA Pharma, Novartis, Sanofi Aventis, Stallergenes, Teva. Grants for clinical studies: Allakos, AstraZeneca, F. Hoffmann-La Roche, GlaxoSmithKline, Otonomy, Strekin; SSe: Employed by Change Accelerator in Respiratory Disease; WJF: Grants from Meda, Allergy Therapeutics. GSK and ALK. LB, LOC, ABS, ASC, JC, LD, SD, ADK, JR, GR, PSJ, GS, PS, AS, UW: no conflict of interest to report.

Authorship contribution

PWH, GS and WJF have made the draft of the algorithm and manuscript, with active participation and input in the discussion and finetuning of the algorithm and manuscript by all experts listed as co-author.

Acknowledgement

We thank Dr Ruth B. Murray (Medscript Ltd) for editorial assistance.

References

1. Hellings PW, Borrelli D, Pietikainen S, Agache I, Akdis C, Bachert C, et al. European Summit on the Prevention and Self-Management of Chronic Respiratory Diseases: report of the European Union Parliament Summit (29 March 2017). *Clin Transl Allergy*. 2017;7:49.
2. Samoliński B, Fronczak A, Kuna P, Akdis CA, Anto JM, Białoszewski AZ, et al. Prevention and control of childhood asthma and allergy in the EU from the public health point of view: Polish Presidency of the European Union. *Allergy*. 2012;67:726–31.
3. Maurer M, Zuberbier T. Undertreatment of rhinitis symptoms in Europe: findings from a cross-sectional questionnaire survey. *Allergy*. 2007;62:1057–63.
4. Tan R, Cvetkovski B, Kritikos V, Price D, Yan K, Smith P, et al. Identifying the hidden burden of allergic rhinitis (AR) in community pharmacy: a global phenomenon. *Asthma Res Pract*. 2017;3:8.
5. Price DB, Scadding G, Bachert C, Saleh H, Nasser S, Carter V, et al. UK prescribing practices as proxy markers of unmet need in allergic rhinitis: a retrospective observational study. *NPJ Prim Care Respir Med*. 2016;26:16033.
6. Hellings PW, Fokkens WJ, Bachert C, Akdis CA, Bieber T, Agache I, et al. Positioning the principles of precision medicine in care pathways for allergic rhinitis and chronic rhinosinusitis - A EUFOREA-ARIA-EPOS-AIRWAYS ICP statement. *Allergy*. 2017;72:1297–305.
7. Canonica GW, Bachert C, Hellings P, Ryan D, Valovirta E, Wickman M, et al. Allergen Immunotherapy (AIT): a prototype of Precision Medicine. *World Allergy Organ J*. 2015;8:31.
8. Hellings PW, Fokkens WJ, Akdis C, Bachert C, Cingi C, Dietz de Loos D, et al. Uncontrolled allergic rhinitis and chronic rhinosinusitis: where do we stand today? *Allergy*. 2013;68:1–7.
9. Scadding GK, Kariyawasam HH, Scadding G, Mirakian R, Buckley RJ, Dixon T, et al. BSACI guideline for the diagnosis and management of allergic and non-allergic rhinitis (Revised Edition 2017; First edition 2007). *Clin Exp Allergy*. 2017;47:856–89.

P.W. Hellings
KU Leuven
Department of Microbiology Immunology and Transplantation
Laboratory of Allergy and Clinical Immunology Research Group
Leuven
Belgium

Tel: +32 16 332211
E-mail: Peter.Hellings@me.com