Allergic rhinoconjunctivitis

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 - NPRANG June 2023



Aim of Presentation

- Overview of the disease
- Common airborne allergens
- Approaches to allergen avoidance
- Medical management of rhinitis and rhino-conjunctivitis
- Immunotherapy

Rhino-conjunctivitis

- Chronic, mostly eosinophilic, inflammation of the nasal mucosa and conjunctiva
- Affects up to 25% of population
- Symptoms include itching, sneezing, watery nasal discharge, and nasal congestion
- Associated ocular symptoms (watery, red and/or itchy eyes)
- Seasonal and/or perennial
- Intermittent or persistent
- Mild, moderate or severe according to the impact on the quality of life





Rhinitis Statistics

- Allergic rhinitis affects between 10% and 30% of all adults and as many as 40% of children
- Up to 57% of adult patients and up to 88% of children with AR have sleep problems leading to daytime fatigue and somnolence, and decreased cognitive functioning (Pawankar R, et al, 2013)

- Associated with considerable loss of productivity and quality of life
- Leads to impaired performance, sleep disturbance, learning ability, drop in grades
- Exacerbates asthma and is a major factor in asthma development (Scadding, 2015)
- Rhinitis is present in up to 95% of patients with asthma (Small 2018)
- Can occur in children as young as 1.4 years of age (Ludman, 2016)
- History, examination and testing cornerstone of diagnosis
- OTC medicines often first line



One airway One disease



Upper and lower airway symptoms and disease are both presentation of a single inflammatory process within the respiratory tract





Common triggers

- House Dust Mite
- Grass
- Tree
- Weeds
- Mould and fungal spores
- Animal dander
- Occupational e.g. flour
- Latex
- Nonspecific triggers such as smoke, dust, viral infections, strong odours, and cold air

Rhinitis/rhinoconjunctivitis

Allergic rhinoconjunctivitis

- Symptoms (nasal itch/sneeze, watery discharge) on allergen exposure
- Conjunctivitis often associated with rhinitis symptoms
- Positive skin prick test or serum-specific IgE to allergens that are relevant according to the history

Infectious rhinitis

- Usually secondary to a viral infection
- Conjunctivitis may be associated with rhinitis symptoms

Nonallergic, non infectious rhinitis • Structural

- Neurogenic
- Hormonal
- Drug induced
 - Irritant
- Other

Different forms of rhinitis may co-exist and may alter the clinical presentation and prevent optimal treatment response

AIT is only indicated for allergic rhinitis /rhinoconjunctivitis, not for other forms of rhinitis

Diagnosis

- SPT
- Specific IgE via hospital lab
- ISAC
- Home allergy tests (ALEX) 294 allergens
- Component testing confirms cross reaction between pollen and food



Guidelines

BSACI updated 2017

Allergic Rhinitis and its Impact on Asthma updated (ARIA 2020) SEL Integrated Guideline



ARIA aims to educate and implement evidence-based management of allergic rhinitis in conjunction with asthma worldwide



Most patients with rhinitis and asthma consult their GP or Pharmacist first

South East London (SEL) Integrated Guideline for the Management of Allergic Rhinitis (AR)



 $\sqrt{\rm Allergic}$ rhinitis is common in children and adults and is a significant cause of morbidity.

 $\sqrt{}$ Symptoms can affect quality of life (2), school performance (3) and impact on family life (4).

V Patients must be evaluated for asthma symptoms. 75% of children with asthma suffer from AR (5) and AR increases the risk of hospitalisation in children with asthma (6)

 $\sqrt{}$ Patients must be asked about eczema and pollen food syndrome.

V Patients must demonstrate their nasal spray technique regularly and adherence to therapy should be established before stepping up therapy.

ARIA Classification of Allergic Rhinitis⁷ Diagnosis 1. Classic symptoms: Rhinorrhoea, pruritus (nose, Persistent throat, mouth), nasal congestion (mouth Intermittent Symptoms **Symptoms** breathing, snoring), sneezing <4 days per week >4 days per week **2. Careful history** (may identify allergic trigger) or <4 consecutive weeks and >4 consecutive weeks 3. Examination of the nose to rule out any structural problems Moderate – Severe Mild one or more of: all of the following: 1) Disturbed sleep 1) Normal sleep 2) Impairment of 2) No impairment of daily activities daily activities 3) Impairment of 3) No impairment of work/school work/school 4) Troublesome 4) Symptoms present Symptoms but not troublesome Approved: June 2019 Review date: June 2021 (or sooner if evidence or practice changes) Not to be used for commercial or marketing purposes. Strictly for use within the NHS

South East London Area Prescribing Committee. A partnership between NHS organisations in South East London: Bexley/ Bromley/ Greenwich/ Lambeth/ Lewisham & Southwark Clinical Commissioning Groups (CCGs) & GSTFT/KCH/SLAM/Oxleas NHS Foundation Trusts and Lewisham & Greenwich NHS Trust

Step u	p treatment if uncontr	olled	ng treatment	Step down when pa	tient gains control
Step 1 If moderate to evere symptoms (ARIA criteria)	Step 2 Start with antihistamine if pruritus dominant or nasal corticosteroid if congestion dominant For information for management in pregnancy and breastfeeding refer to <u>CKS</u>		Step 3	Step 4 (Primary care or specialist initiation)	Step 5 Specialist allergy clinic
Allergen avoidance easonal allergic hinitis – please ee <u>Allergy UK</u> uidance and <u>SACI guidance</u> nd <u>IHS Choices</u> erennial (House ust mite) – lease see <u>BSACI</u> uidance	Regular long acting non-sedating antihistamine (see p3 for OTC options in adult patients) See BNF and BNFC for current dosing for age group and formulations available <u>1st line:</u> Cetirizine (from age 1 year. Use twice daily regimen in <12 years) or Loratadine (from age 2 years) <u>2nd line:</u> (can consider if trial of above fails): Fexofenadine (from age 6 years) Regular nasal antihistamine <u>Azelastine</u> Only if oral antihistamines are not appropriate	Regular nasal corticosteroid spray* (see p3 for OTC options in adult patients) Demonstrate & check technique See <u>BNF</u> and <u>BNFC</u> for current dosing for age Mometasone furoate (50micrograms per spray) From age 6 years Or Fluticasone <u>furoate</u> (Avamys®) (27.5 micrograms per spray) From age 6 years. Good effect on eye symptoms Or Fluticasone <u>propionate</u> (e.g. Flixonase®) (50 micrograms per spray) From age 4 years For patients taking cobicistat or ritonavir use: Beclometasone dipropionate (50 micrograms per spray) See MHRA advice Dec 2016	Trial of oral antihistamine and nasal corticosteroid as per products in Step 2	Regular nasal antihistamine, nasal corticosteroid and oral antihistamine Consider switching separate nasal antihistamine and nasal steroid spray (especially if patient is already on fluticasone propionate nasal spray) to: Fluticasone propionate with azelastine spray (Dymista®) From age 12 years Good effect on eye symptoms.	Specialist initiation only Leukotriene receptor antagonis: (e.g. montelukast) Can be considered in asthmatic patients See <u>BNF</u> and <u>BNFC</u> for current dosing for age Specialist initiation and continued prescribing only Allergen specific immunotherapy South East London Area Prescribing Committee Approved: June 2019 Review date: June 2021 (or sponer if evidence or practice

SEL Integrated Guideline for the Management of Allergic Rhinitis (AR)

Paediatrics	Adults	ENT Red Flags for urgent referral		
Top Tips 1. For seasonal rhinitis, start nasal spray 1-2 weeks before onset of appropriate pollen season 2. Nasal steroids unlikely to work if there is nasal blockage due to secretions. Try nasal steroid drops or pre-dosing with topical decongestant for 5 days 3. Avoid sedating antihistamines, and intranasal beclometasone (e.g. Beconase®) as it can have systemic effects due to a high bioavailability (due to interactions beclomethasone is the preferred product in those taking cobicistat or ritonavir however. See MHRA advice Dec 2016)	For seasonal rhinitis, start nasal spray 1-2 weeks before onset of appropriate pollen season If eye symptoms present consider: •Olopatadine eye drops (see APC recommendation) •Sodium cromoglicate eye drops In severe cases of nasal obstruction thought to be due to allergic rhinitis a short course (e.g. 5 days) of prednisolone 0.5mg/kg could be considered (adults only, max 2 courses per year) AVOID: •Sedating antihistamines •Depot corticosteroids •Chronic use of decongestants or nasal beclometasone, as has high bioavailability	 Unilateral symptoms including blockage, clear rhinorrhoea and facial pain Serosanguinous discharge Visual and neurological signs (considering sinonasal malignancy) Failure of 3 months maximum medical therapy, particularly where nasal blockage and anosmia remain significant symptoms 		
 4. Avoid and chronic use of decongestants 5. If eye symptoms present consider: Olopatadine eye drops (from age 3) (see APC recommendation) Sodium cromoglicate eye drops 	 The following may be an indication for referral to Allergy Specialist Inadequate control of symptoms on conventional treatment 1.Allergen/trigger identification 2.Consideration of desensitisation 3.Recurrent nasal polyps 4.Multisystem allergy (e.g. rhinitis with asthma, eczema or food allergy) 5.Occupational rhinitis 	References1.Ant K et al. J. Allergy 2009;64:123–148.2.Silva CHM et al. Braz J Otorhinolaryngol2009;75:642–649.3.Walker S et al. J Allergy Clin Immunol2007;120:381–387.4.Emin O et al. Int J Pediatr Otorhinolaryngol		
 to Paediatric Allergy Specialist 1. Children with AR who are unresponsive and/or intolerant to conventional treatment 2. Children with diagnostic uncertainty and in whom further investigations (skin prick test +/- slgE) would be helpful 4. Children who may be considered for 	 For adults, the following are available OTC without prescription, which patients could consider buying: Fluticasone propionate 50mcg nasal spray Beclometasone 50mcg nasal spray (not preferred for routine prescribing, though may be cheaper than fluticasone OTC. Preferred product for patients on cobicistat or ritonavir, see MHRA advice Dec 2016) Loratadine tabs, liquid 	2009;73:1795–1798 5.Ballardini N et al. Allergy 2012;67:537–544. 6.Lasmar L et al. J Pediatr 2007;83:555–561. 7.Bousquet J et al. Allergy 2008;63:8-160 8.Roberts G et al. Allergy 2013; 68: 1102–1116 9.Scadding G K et al. Clinical & Experimental Allergy 2008, 38: 19–42. Approved: June 2019		
desensitisation 5. Multisystem allergy (rhinitis with eczema, asthma or food allergy)	 Cetirizine tabs, liquid Sodium cromoglicate eye drops Xylometazoline & antazoline eye drops (Otrivine Antistin[®]) 	Approved: June 2019 (or sooner if evidence or practice changes) Not to be used for commercial or marketing purposes. Strictly for use within the NHS		

Management

The aims of management are to control symptoms and reduce inflammation



Despite medication, a significant number of patients continue to experience symptoms that impair their quality of life



Avoidance is recommended in all guidelines

Adherence is often poor

- 25% of all asthma and allergy patients do not collect their prescriptions
- Non-adherence may be non-intentional or intentional
- Bedrock of adherence and positive healthcare behaviours is an acceptance of the condition
- Doubts over the need for medication and avoidance measures are closely linked to poor adherence and risk taking behaviour
- Burden of care, interventions and regimens can be difficult to follow
- Asthma and allergy patients bargain with themselves



Avoidance of Pollens

Pollen calendar

The timing and severity of pollen seasons differ across the UK from south to north and east to west. This pollen calender shows the general timing that the main allergenic plants are in flower.

Pollen season: Plant type Feb Sep Aug lan Mar Apr May lun lul Alder Hazel Yew Elm Willow Tree Poplar Birch Ash Plane Oak Pine Lime Grass Oil seed rape Plantain Weed Nettle Dock

Information sourced from Univercity of Worcester www.worcester.ac.uk

Mugwort



Thirty years of supporting people living with allergy

Pollen Avoidance

- Watch the pollen count, know the relevant pollen season
- Use an air filter
- Keep windows closed
- Wraparound sunglasses, Vaseline
- Shower when you get home, wash pollen out of hair and change clothes
- Don't dry clothes on the line if the pollen count is high



Avoidance of HDM



Evidence of avoidance measures

- Cochrane review 2015 no evidence to support HDM avoidance for eczema
- Mite-impermeable encasings are effective in reducing the number of mite-sensitized children with asthma attending the hospital with asthma exacerbations but not the number requiring oral prednisolone (2017)



Non-pharmacological Management

- Nasal saline douching Cochrane review 2018
- Salt nasal sprays all ages (cheap and safe)
- Allergy products containing Ectoin (protects mucous membranes in eye and nose) can offer relief
- Airsonett laminar airflow device
- Complementary and Alternative









Antihistamines

- Histamine-1 receptor antagonist
- Competes with histamine for H1 receptor sites
- BUT doesn't displace histamine that is already bound to receptors
- First generation Chlorphenamine and Diphenhydramine
- Second generation of AH only minimally penetrate BBB
- Cetirizine, Levocetirizine, Loratadine, Desoloratadine, Fexofenadine are recommended for SAR management

Side effects and interactions

- May increase sedative and respiratory depressant effects of CNS depressants e.g. alcohol and tranquilizers
- Loratadine may cause serious cardiac effects when taken with macrolide antibiotics, fluconazole, ciprofloxacin and clarithromycin
 - Side effects of CNS include dizziness, fatigue, disturbed coordination, muscle weakness
 - GI reactions include loss of appetite, nausea and vomiting, constipation, diarrhea, dry mouth nose or throat



Cardiac reactions include hypo/hypertension, rapid pulse, arrhythmias



Caution with epilepsy, renal failure and fruit juices

Topical antihistamine

Azelastine hydrochloride Rhinolast (140mcg per spray)

6yrs and above one spray twice daily

Optilast 0.05% (500mcg per 1 ml)

twice daily up to 4 times per day

Mast cell stabilizers

Delayed results

Fewer side-effects than the other treatments

Lodoxamide >4yrs up to 4 times per day into each eye for up to 4 weeks

Sodium Cromoglicate up to 4 times per day into each eye

Only offer symptom relief but can sting when administered

Leukotriene Receptor Antagonists



Caution in hepatic impairment and has several potential drug interactions

Leukotriene Receptor Agonists Interactions and ADRs



Increased toxicity with erythromycin, fluconazole, metronidazole



Decreased effectiveness with phenytoin, phenobarbital



GI (diarrhoea, nausea, vomiting), headaches, fever, skin reactions



Sleep disturbances, dizziness, drowsiness, irritability, cough

Intra nasal corticosteroids

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Anti-inflammatory action via gene regulation of immune cells



Inhibit the production of cytokines, leukotrienes, prostaglandin, reduces the recruitment of eosinophils and release of inflammatory mediators



Image: bronchial inflammation
 Image: bronc



Nasal Sprays: Choices and doses

- Fluticasone (Avamys/Nasofan/Flixonase/Pirinase)
 4-11yrs 50mcg once or twice daily
 12 yrs and above 100mcg once or twice daily
- Mometasone (Nasonex/Clarinaze)

3-11 yrs 50mcg daily

12 yrs and above 100-200mcg daily

• Beclometasone (Beconase/Nasobec)

6 yrs and above 100mcg twice daily (increased bio-availability)

Budesonide (Rhinocort)

6 yrs and above 64mcg twice daily

- Combination Fluticasone and Azelastine (Dymista) 12 yrs and above one spray twice daily
- Combination Olopatadine and Mometasone (Ryaltris)

12 yrs and above two sprays twice daily

• Azelastine (Rhinolast) – Antihistamine only

Over 4 years – Twice to four times daily

• Sea Water Spray (Sterimar)

3 yrs and above two – six times daily

How to use a nasal spray

Nurses in Allergy Standard Operating Procedure



Topical Nasal Corticosteroid Spray

Using a topical nasal corticosteroid spray is recognised as a first line treatment to control nasal congestion for both allergic and non-allergic rhinitis. This type of spray is often referred to simply as a steroid nasal spray.

Steroid nasal sprays are used for both persistent and seasonal rhinitis. Rhinitis caused by perennial allergens such as the house dust mite are more likely to cause persistent symptoms and require continuous long-term treatment especially when a patient has symptoms such as nasal blockage (1,2).

Systemic absorption of nasal corticosteroid sprays depend on the bioavailability of the drug. Long term use of corticosteroid nasal sprays are considered relatively safe, but it is advisable to use a spray with a low systemic bioavailability when patients require continuous treatment for extended periods (3).

The application of a steroid nasal spray is localised to the affected area. The spray works by reducing inflammation and associated symptoms of increased mucus production and possibly sneezing. It does not work immediately and can take up to two weeks before a patient perceives the benefit from using a steroid nasal spray. In seasonal altergic hinitis (haydvery treatment should begin two weeks before symptoms are expected to start (4, 5) therefore ensuring the efficacy of the spray by the time the trigger altergen is in the air.

It has been shown that nasal douching before the use of a steroid nasal spray will enhance efficacy and generally improve symptomatic control (2).

Nasal Corticosteroid Sprays							
Generic Name	Proprietary Name	Can be prescribed for	Bioavailability				
Triamcinolone	Nasocort	over 12 years					
Acetonide		two sprays each nostril od	46%				
		children 6-11 years					
		one spray each nostril od Up to bd					
		children 2-6 years					
		one spray each nostril od					
Beclometasone	eclometasone Beconase over 6 yrs		4.4%				
Dipropionate		two sprays each nostril bd	4470				
Budesonide	Rhinocort Aqua	over 12 years					
		two sprays each nostril bd	31%				
Flunisolide	Syntaris	over 14 years					
		two sprays each nostril bd	20-30%				
		children 5-14 years					
Fluticasone &	Dymista	over 12 years	>				
Azelastine	Dynnota	one spray per nostril bd	1.86%				
Mometacone Euroate	Naconey	over 12 years	1.0070				
Montetasone i dioate	Nasonex	two sprays each nostril od	0.46%				
		children 6-11 years					
		one spray each nostril od Up to bd					
Fluticasone Propionate	Flixonase, Nasofan	over 12 years					
Fluticasone Furoate	Avamys	two sprays each nostril od	0.42%				
		children 4-11 years					
		one spray each nostril od Up to bo	1				
		Avamys - from 6 years					

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Olopatadine / ometasone Furoat

Ryaltris

Decongestants

Oral or nasal decongestants (pseudoephedrine, phenylephrine)

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Relieve nasal congestion but not recommended as part of rhinitis guidelines



Contraindicated in uncontrolled hypertension, severe coronary artery disease

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Prolonged use leads to rhinitis medicamentosa (only use for 3-5 days)



Side effects - agitation, insomnia, headache, palpitations limit use

Ocular Symptom Control



Avoidance, cold compresses and eyewashes

Oral antihistamines

Mast cell stabilizers (Sodium Cromoglycate)

Olopatadine (Epinastine and mast cell stabilizer >3yrs twice daily

Ketofall (Ketotifen) >3yrs twice daily

Topical Corticosteroids

Refer to Opthalmologist for more advanced treatment e.g. Verkazia (Ciclosporin from > 4 years)

Early phase allergic conjunctivitis



Both seasonal and perennial allergic conjunctivitis are two acute disorders

Inflammation of the conjunctiva (the membrane covering the white part of the eye)



Increased vasodilation and lacrimation



Photophobia

E Sensation of foreign body





Late Phase allergic conjunctivitis



Chronic allergic disease

eosinophils, conjunctival fibroblasts, epithelial cells, mast cells and TH2 lymphocytes Vernal keratoconjunctivitis

graded based on severity and common in summer chronic, bilateral disease with no seasonal correlation

Atopic

keratoconjunctivitis



Immunotherapy



How does IT work?

- Promotes induction of blocking protective IgG antibodies
- Increases regulatory T lymphocytes
- Shifts allergen specific antibody production from IgE to IgG4
- Blunts seasonal increases in IgE levels
- Downregulates IgE mediated histamine release
- Immune deviation from TH2 to TH1 allergen specific response
- Protects against disease progression
- Symptomatic improvement for years after the treatment is discontinued



Side effects

SCIT

 Localized swelling and itching, lethargy, rhinitis symptoms, higher risk of severe reactions with SCIT especially venom

SLIT

 Oral pruritis, ulcers, lip and oral swelling, itchy ears, sneezing, throat irritation, abdominal pain, nausea, can induce rhinitis symptoms, very rarely severe

Criteria for Immunotherapy



Crank

Symptoms strongly suggestive of AR, with or without conjunctivitis



Evidence of IgE sensitization (positive SPT and/or serum-specific IgE) to one or more clinically relevant allergen



Experience moderate-to-severe symptoms which interfere with usual daily activities or sleep despite regular and appropriate pharmacotherapy and/or avoidance strategies





Take advantage of long-term effects on AR and it's potential to prevent asthma



Pollen Food Syndrome

- Birch apple, pear, cherry, peach, nectarine, apricot, plum, kiwi, hazelnut, almond, celery, carrot, potato
- Birch/Mugwort celery, carrot, spices, sunflower seed, honey
- Grass melon, watermelon, orange, tomato, potato, peanut
- Ragweed Melon, banana, courgette, cucumber
- Plane hazelnut, peach, apple, melon, kiwi, peanuts, maize, chickpea, lettuce, green beans

Summary



Allergy to airborne antigen is extremely common and triggers allergic rhinitis and asthma

It should be treated as one airway one disease

Allergen avoidance can be effective in specific situations but is hard to achieve and often expensive and impractical

Daily non-drowsy antihistamines and steroid nasal spray are the main daily recommended treatments

Immunotherapy is the only disease modifying option