

#### Derbyshire Medicines Management, Prescribing and Guidelines DERBYSHIRE PRIMARY CARE FORMULARY

#### Chapter 3: RESPIRATORY SYSTEM Updated: March 2024

### The following prescribing guidelines are relevant to the respiratory chapter and can be found here

- Children's referral guideline for SLIT- Grazax and Acarizax (Derby Children's Hospital only)
- Anaphylaxis treatment for adults and children
- Asthma management in adults/ Asthma management in children
- Chronic Obstructive Pulmonary Disease guideline
- Greener Inhaler Prescribing guidance
- Oxygen guideline

Relevant Resources

- COPD detailing aid
- Greener inhaler choice: Flowchart/ Be Greener and Breathe Better patient information leaflet
- Stepping-down combination asthma inhaler therapy: Adults over 17 years of age
- Optimisation of inhaled corticosteroid (ICS) in COPD
- Adult Asthma Diagnosis Algorithm <u>North Derbyshire</u>, <u>South Derbyshire</u>
- Respiratory Action Plan and Asthma Self-management Plan
- <u>RightBreathe</u> inhaler information/training videos
- Inhaled corticosteroid equivalent doses

# Reducing the carbon impact of inhalers

Metered dose inhalers (MDIs), including breath-actuated MDIs, contain propellants known as hydrofluorocarbons (HFCs) which are powerful greenhouse gases and can contribute to global warming. Dry powder inhalers (DPIs) and soft mist inhalers (SMIs) such as a Respimat do not contain propellant so they have a lower carbon footprint than other inhalers.

NHS Derby and Derbyshire ICB/System partners support the prescribing of inhalers with a reduced carbon footprint such as DPIs and SMIs, wherever clinically appropriate and acceptable to the patient. Medication reviews, Structured Medication Reviews or planned Asthma Reviews taking place in primary care should consider moving or facilitating patients to lower carbon options where it is clinically appropriate to do so. See <u>Greener inhaler prescribing guidance</u>.

All formulary dry powder inhalers contain lactose and are contraindicated in patients with hypersensitivity to lactose or milk proteins. Refer to The SPC for full prescribing information

<u>MHRA July 2018</u> Pressurised metered dose inhalers (pMDIs): risk of airway obstruction from aspiration of loose objects. Remind patients to check and remove the mouthpiece cover properly before inhaling a dose and to shake the inhaler to remove loose objects that may have become trapped in the inhaler during storage.

# 3.1 Bronchodilators

### 3.1.1 Adrenoreceptor agonists

Short acting beta agonist (SABA)	Traffic Light Classification	Licenced	Additional information
<ul> <li>Salbutamol preparation</li> <li>100 micrograms CFC Free inhaler (MDI) (Salamol)</li> </ul>	GREEN	Adults and children ≥4 years of age	The metered dose inhaler (MDI) is usually the most cost-effective delivery device for salbutamol and should be considered for new patients and as a matter of routine for
• 100 micrograms Easyhaler (DPI)	GREEN		all paediatric patients. Salamol MDI is the preferred choice salbutamol inhaler due to
200microg Ventolin Accuhaler (DPI)	GREEN		lower carbon footprint compared to other salbutamol MDIs. Patients who are unable
100 micrograms Breath-actuated CFC free inhaler (MDI)	GREEN		to use a standard salbutamol MDI may find a dry powder device or breath-actuated inhaler more acceptable. The brand must

Nebuliser solution 2.5mg/2.5ml			be specified to avoid confusion with these devices. Nebulised bronchodilators should not be prescribed unless a formal nebuliser assessment has been carried out Do not use 5mg salbutamol nebules for COPD, as evidence does not support doses above 2.5mg. Nebulisers should be used with extreme caution in children and only under the care of a respiratory paediatrician (MHRA Aug 2022).
Long acting beta agonist (LABA)	Traffic Light Classification	Licenced	Additional information
<ul> <li>Formoterol preparations</li> <li>12 micrograms Easyhaler (DPI)</li> </ul>	GREEN	Adults and children ≥6 years of age	First-line LABA
12 micrograms CFC free inhaler (MDI) (Atimos Modulite)	GREEN	Adults and children ≥12 years of age	Alternative first-line LABA for patients requiring an MDI
<ul> <li>Salmeterol preparations</li> <li>25 micrograms CFC Free inhaler (MDI) (Soltel)</li> </ul>	GREEN 2 <sup>nd</sup> line LABA	Soltel - Adults and children >12 years of age Serevent - Adults and children ≥4 years of age	<b>Soltel</b> is currently the formulary choice N.B. All brands apart from Serevent contain soya lecithin – <b>contra-indicated</b> <b>in peanut or soya allergy</b> ). If the patient has a soya or peanut allergy, then prescribe as the brand Serevent

### 3.1.2 Antimuscarinic bronchodilators

Short acting antimuscarinic bronchodilators (SAMA)	Traffic Light Classification	Licenced	Additional information
<ul> <li>Ipratropium</li> <li>20 microg inhaler CFC free (MDI)</li> </ul>	GREEN	COPD and asthma Adults and children (including those <6 years of age)	Asthma – MDI is not recommended by SIGN/BTS or NICE for routine asthma management. Nebulised solution may be used as an add-on treatment for a severe asthma attack.

- 1. Ipratropium nebulizer solution for COPD should only be used in severe COPD patients [FEV1<30%] after initiation by a specialist.
- Long acting muscarinic antagonists (LAMAs) monotherapy are no longer recommended first line in the management of COPD- see <u>local guidance</u>. When initiating a LAMA, take care to ensure the SAMA is stopped.
- 3. <u>Tiotropium</u> is the preferred LAMA for existing stable patients on single component LAMA treatment.
  - 18 microgram inhalation powder (DPI) (Tiogiva)- Advise patients NEVER to insert the capsule directly into the mouthpiece, always follow the instructions provided with the inhaler. Tiogiva inhaler device should be replaced every 6 months.
  - 2.5microgram/ inhalation solution cartridge CFC Free (soft mist inhaler) (Spiriva Respimat)-Respimat inhalers are re-usable device with cartridge. Each re-usable inhaler may be used with up to six cartridges. It is **Grey** after consultant/specialist initiation for asthma.
- 4. Tiotropium
  - <u>MHRA Feb 2015</u> prescribers should take the risk of cardiovascular side effects into account when prescribing inhaled tiotropium to patients with certain cardiac conditions, who were excluded from clinical trials of tiotropium (See SPC for detail).
  - Plasma concentration of tiotropium increases with decreased renal function in patients with creatinine clearance ≤ 50 ml/min- only use if the expected benefit outweighs the potential risk.
- 5. Alternative LAMA includes
  - Glycopyrronium bromide inhaler 44microgram caps (DPI) (Seebri Breezhaler)- Once daily LAMA (2nd line). Only use if the expected benefit outweighs the potential risk in patients with severe renal impairment (eGFR below 30 ml/min/1.73 m<sup>2</sup>), including those with end-stage renal disease requiring dialysis. These patients should be monitored closely for potential adverse reactions.
  - Umeclidinium inhaler 55microgram/ inhalation (DPI) (Incruse Ellipta)- once daily LAMA (2nd line)
  - Aclidinium inhaler 322microgram/ inhalation (DPI) (Eklira Genuair)- twice daily LAMA (3<sup>rd</sup> line). (Each delivered dose contains 375mcg of aclidinium bromide, equivalent to 322mcg.)

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Combination inhalers N.B. Prescribe by brand

LABA/LAMA combinations	Traffic Light Classification	Licenced	Additional information
Tiotropium and olodaterol (Spiolto Respimat) (SMI)	GREEN 1 <sup>st</sup> line LABA /LAMA	COPD	Respimat inhalers are re-usable device with cartridge. Each re-usable inhaler may be used with up to six cartridges.
Glycopyrronium and Indacaterol (Ultibro Breezhaler) (DPI)	GREEN 1 <sup>st</sup> line LABA /LAMA	COPD	Choice should be based on patient tolerance, ease of use, and
Umeclidinium and vilanterol (Anoro Ellipta) (DPI)	GREEN 1 <sup>st</sup> line LABA /LAMA	COPD	environmental impact of the inhaler device.
Aclidinium and formoterol (Duaklir Genuair) (DPI)	GREEN 1 <sup>st</sup> line LABA /LAMA	COPD	LABA/LAMA combination inhaler is recommended for COPD patients who remain breathless or have exacerbations
Glycopyrronium + formoterol (Bevespi Aerosphere) (MDI)	<b>GREEN</b> 1 <sup>st</sup> line LABA/LAMA for patient requiring an MDI	COPD	despite SABA or SAMA treatment and present with no asthmatic features or features suggestive of steroid responsiveness.

 <u>Roflumilast</u> is **GREY**- specialist initiation. Roflumilast is a phosphodiesterase type-4 inhibitor with antiinflammatory properties. It is used as an add-on to bronchodilator therapy in adults with severe COPD with chronic bronchitis as per NICE TA461. Ongoing GP prescribing and care of patients on roflumilast should only be considered if patient is stable and free from adverse reactions, after a minimum of 3 months roflumilast treatment under the respiratory specialist. For more details see local <u>COPD guideline</u>.

# 3.1.3 Theophylline

Theophylline SR tablets (Uniphyllin Continus) 200mg, 300mg, 400mg

- Common interactions for theophylline which increase clearance and it may therefore be necessary to increase dosage to ensure therapeutic effect include barbiturates, carbamazepine, lithium, phenytoin, rifampicin, primidone, ritonavir. (See SPC for full details) The following reduce clearance and a reduced dosage may therefore be necessary to avoid sideeffects: allopurinol, cimetidine, corticosteroids, diltiazem, macrolide antibiotics (e.g., erythromycin), frusemide, and oral contraceptives. (See SPC for full details)
- 2. Smoking can increase theophylline clearance and increased doses of theophylline are therefore required; dose adjustments are likely to be necessary if smoking started or stopped during treatment.

### 3.1.5 Peak flow meters, inhaler devices and nebulisers

Appropriate peak flow meter Appropriate spacer device	EasyChamber Spacer fits most MDIs
	Volumatic

### See Right Breathe website for information on spacer compatibility.

- 1. EasyChamber fits most MDIs, and is compatible with formulary choices. Although not listed, this includes Luforbec.
- 2. Follow usage and cleaning instructions supplied with the spacer.
- 3. Spacers should not be regarded as interchangeable patients whose asthma is well-controlled and who are using a spacer should always use the same type of spacer and not switch between spacers. Different spacers may deliver different amounts of inhaled corticosteroid, which may have implications for both safety and efficacy see <u>Drug Safety Update</u>, July 2008
- 4. All patients with an acute asthma attack at home, and symptoms not controlled by up to 10 puffs of salbutamol via a pMDI and spacer, should seek urgent medical attention. (BTS/SIGN 2019)
- 5. Standard-range EU peak flow meters are suitable for both adults and children; low-range peak flow meters are appropriate for severely restricted airflow in adults and children.
- 6. <u>MHRA Aug 2022</u> : home use of nebulisers in paediatric asthma should be initiated and managed only by specialists (under a treatment plan). Use of a nebuliser purchased independently of medical advice for use in the home to deliver nebulised asthma rescue medications to children can mask a deterioration in the underlying disease and may increase the risk of potentially fatal delays in seeking medical attention if asthma deteriorates.

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### 3.2 Corticosteroids

**N.B. Prescribe by brand**; use MDI with spacer device for all doses of inhaled corticosteroid. **See <u>adults</u> and <u>children's</u> asthma guidance** for inhaled corticosteroid doses

Corticosteroids (ICS)	Traffic Light Classification	Licenced	Additional information
<ul> <li>Beclometasone dipropionate</li> <li>Soprobec MDI 50, 100, 200, 250 micrograms</li> </ul>	GREEN	Asthma- adults and children	BNFC doses for Soprobec in asthma: Child: 100microg BD increased if necessary up to 400microg daily in 2-4 divided doses.
• Kelhale MDI 50,100 microgs (extra fine particles)	GREEN	Asthma- adults >18 years	Kelhale is therapeutically equivalent to Qvar.
QVAR MDI 50, 100     micrograms(extra fine     particles)	GREEN	Asthma- ≥5 years of age	SPC doses for Qvar in asthma- Children aged 5 years and over: Mild – 50 micrograms BD Moderate – 50 to 100 micrograms BD Severe- 100 micrograms BD Kelhale/Qvar or fluticasone are twice as potent as Soprobec (e.g. Kelhale/Qvar and fluticasone 50 microg is equivalent to Soprobec100microgs)
<ul> <li>Budesonide</li> <li>Easyhaler (DPI) 100, 200,400 microg/ inhalation</li> </ul>	GREEN	Asthma- ≥6 years of age	dose equivalent to standard particle beclomethasone (Soprobec/ Clenil)
<ul> <li>Fluticasone</li> <li>Flixotide Accuhaler (DPI)100 microgram</li> <li>Flixotide Evohaler (MDI) 50 microgram</li> </ul>	GREEN for children GREY for adults	Asthma- >4 years of age	As per <u>children's</u> asthma guidance Kelhale/Qvar or fluticasone are twice as potent as Soprobec (e.g. Kelhale/Qvar and fluticasone 50 microg is equivalent to Soprobec100microgs)

Combination inhalers N.B. Prescribe by brand. See asthma and COPD guideline.

LABA/ICS combination	Traffic Light Classification	Licenced	Additional information
Budesonide and formoterol			
• Fobumix easyhaler DPI 80/4.5, 160/4.5, 320/9	<b>GREEN</b> 1 <sup>st</sup> line LABA/ICS combination inhaler	Asthma (≥18 years of age) & COPD	Fobumix 80/4.5, 160/4.5, 320/9 is equivalent to Symbicort 100/6, 200/6, 400/12 respectively Fobumix 80/4.5 licensed for asthma only.
• WockAIR DPI 160/4.5, 320/9	GREEN	Asthma (≥12years of age) & COPD	WockAIR and DuoResp 160/4.5, 320/9 is equivalent to Symbicort 200/6, 400/12 respectively. WockAIR is the most cost effective
<ul> <li>DuoResp Spiromax DPI 160/4.5, 320/9</li> </ul>	GREEN	Asthma (≥12 years of age) & COPD (≥18 years of age)	DPI where appropriate but is only available in two strengths.
Symbicort Turbohaler DPI     100/6	GREEN	Asthma (≥6 years of age)	
Symbicort Turbohaler DPI 200/6, 400/12	GREEN	Asthma (≥12 years of age) & COPD	Symbicort pMDI contains propellant HFA227ea which has a significantly higher carbon footprint than other propellants and
Symbicort MDI 200/6	GREEN	COPD	so should be avoided where possible.
Beclometasone (extra fine) and			
formoterol		Asthma (≥18 years	
Luforbec MDI 100/6	GREEN 1 <sup>st</sup> line	5,	100 micrograms of beclometasone
Luforbec MDI 200/6	for patients	Asthma (≥18 years	dipropionate via Luforbec/Fostair products
	requiring an MDI	of age)	are equivalent to 250 micrograms of beclometasone dipropionate in standard
Fostair Nexthaler DPI 100/6	GREEN	Asthma (≥18 years	particle CFC-free inhalers.
		of age) & COPD	
Fostair Nexthaler DPI 200/6		Asthma (≥18 years of age)	Inhaled corticosteroid equivalent doses

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Fluticasone propionate and			See children's asthma guidance.
salmeterol			See children's astillia <u>guidance</u> .
<ul> <li>Combisal MDI 50/25</li> <li>Seretide MDI 50</li> <li>Seretide Accuhaler DPI 100</li> <li>Fixkoh Airmaster DPI 50/100</li> <li>Combisal MDI 125/25, 250/25</li> <li>Fixkoh DPI 500/50</li> </ul>	GREEN for children GREY for adults GREY - limited place of therapy for adults	Asthma ≥4 years of age Asthma ≥12 years of age Asthma ≥12 years of age Asthma >12 years of age & COPD	Combisal MDI is the cost-effective alternative for Seretide evohaler. Children receiving Seretide/combisal 50 MDI or Seretide/ Fixkoh 100 Accuhaler should be reviewed at age 18 year and considered for the formulary choice ICS/LABA combination inhalers. Fluticasone/salmeterol combination has limited place of therapy in adults- following choices in local COPD guidance only.
<ul> <li>Fusacomb Easyhaler DPI 500/50</li> </ul>		Adults for COPD and asthma (≥12 years of age)	Fixkoh DPI/Fusacomb Easyhaler are cost- effective alternative for Seretide Accuhaler 500
Triple combination (ICS+ LABA+ LAMA)	Traffic Light Classification	Licenced	Additional information
Beclometasone, formoterol and glycopyrronium (Trimbow NEXThaler DPI) Fluticasone, vilanterol and umeclidinium (Trelegy) (DPI)	GREY 1 <sup>st</sup> line triple combination GREY alternative 1 <sup>st</sup> line triple combination	COPD	Indicated for the maintenance treatment of moderate to severe COPD. Triple therapy is reserved for exceptional use in severe disease in the presence of persistent exacerbations or patient is still limited by symptoms despite other treatments. Triple therapy in a single inhaler may be preferable for people who have difficulty using
Beclometasone, formoterol and glycopyrronium (Trimbow) (MDI) Budesonide, formoterol and	GREY after consultant/ specialist initiation for asthma. GREY	COPD Asthma in adults (172/5/9 microg licensed for asthma only) COPD	more than one device or who find their medication regimen difficult or confusing and have trouble complying with treatment. However triple therapy lack flexibility and makes it difficult to amend the individual medicines if treatment needs changing for any reason.
glycopyrronium (Trixeo aerosphere) (MDI) Mometasone, indacaterol and glycopyrronium (Enerzair Breezhaler) (DPI)	GREY after consultant/ specialist initiation for asthma.	Asthma in adults	Use of combination product is often more cost effective than using the separate components. Enerzair Breezhaler and Trimbow MDI are licensed for asthma in adults not adequately controlled with a maintenance combination of a LABA+ ICS who experienced one or more asthma exacerbations in the previous year.

- 1. All doses of inhaled steroid delivered via an MDI should be via a spacer device. This form of administration improves delivery of drug to the airways reducing local effects e.g., oral candida and also reduces deposition in the oropharynx (thereby reducing local adverse effects and the amount of systemic absorption).
- 2. SIDE EFFECTS: Inhaled steroids do not usually cause adrenocortical suppression at normal doses (up to about 1500 micrograms/day of beclometasone (standard particle size)). However, in paediatric practice adrenal suppression has been seen at doses as low as 800 micrograms/day (standard particle size). All children on these doses should have careful growth monitoring and a written management plan advising about the risk of adrenal suppression. Oral candidiasis may be related to dose and dose frequency. Patients should be advised to rinse the mouth after use of high dose steroids to help minimize this. Dysphonia can occur. The use of a large volume spacer device (Volumatic) may reduce these local adverse effects.
- 3. Inhaled corticosteroids and adrenal suppression in children 'Adrenal suppression may be underrecognised'. Prescribers are reminded that:
  - It is important to monitor therapy regularly and titrate down to the lowest dose at which effective control of asthma is maintained.
  - Growth (height and weight centile) should be monitored at least annually in children with asthma.

- 4. If a doctor considers that a child's asthma is not controlled and an increase to a high dose of inhaled corticosteroid plus a LABA, the child should be referred to a specialist in the management of paediatric asthma.
- 5. Patients on high dose ICS (>1000 microgram BDP equivalent/day) should be advised to carry a blue steroid treatment card.
- 6. Advise patients to report any blurred vision or other visual disturbances due to rare risk of central serous chorioretinopathy with corticosteroids (MHRA August 2017).
- 7. The role of Single maintenance and reliever therapy (SMART) and Maintenance and reliever therapy (MART) has a limited place for patient therapy and selection. MART is suitable for patients who despite good adherence to regular maintenance doses of a combination ICS/LABA inhaler, have;
  - 1. Inadequate asthma control and are in frequent need of reliever medication.
  - 2. Asthma exacerbations in the past requiring medical intervention.
- 8. National patient safety alert <u>August 2020</u> steroid emergency card to be issued by prescribers to help healthcare staff to identify appropriate patients and gives information on the emergency treatment if they are acutely ill, or experience trauma, surgery or other major stressors. For further guidance on this see <u>when should I issue a steroid emergency alert card.</u>
- Inhaled Budesonide for treatment of COVID-19- Inhaled budesonide should no longer be considered as a treatment for individuals with COVID-19 infection other than within the context of a clinical trial. People already using budesonide for conditions other than COVID-19 should continue treatment if they test positive for COVID-19. See NICE <u>NG191 Covid-19 rapid guideline: managing</u> <u>COVID-19</u> (updated Dec 2021)

### 3.3.2 Leukotriene receptor antagonists

See adult and children's asthma guidance

Montelukast tabs 10mg, chewable tabs 4mg, 5mg, granules 4mg

- Granules are more expensive and should be reserved for children between 6 months and 2 years old or if chewable tablets not suitable. 4mg chewable tablets are licensed in children 2-5 years old; 5mg chewable tablets are licensed in children 6-14 years old; 10mg tablets licensed for adults and children 15 years and over
- MHRA September 2019 Prescribers should be alert for neuropsychiatric reactions in patients taking montelukast and carefully consider the benefits and risks of continuing treatment if they occur. Patients, parents and carers should be warned of possible adverse reactions affecting sleep, behaviour and mood. If these occur the patient, parent or carer should be advised to contact their doctor or nurse for advice on stopping montelukast.

### 3.4.1 Antihistamines

For treatments of minor self-limiting conditions such as mild to moderate hay fever, self-care is encouraged. Treatments are available to purchase over-the-counter.

Cetirizine 10mg tabsNon-sedatingLoratadine 10mg tabs, oral solution 5mg/5mlNon-sedatingChlorphenamine tabs 4mg, oral solution Sugar-Free 2mg/5mlSedating

1. JAPC has classified Alimemazine as **Do Not Prescribe (DNP)** due to lack of cost-effectiveness. Suitable alternative that may be considered is promethazine (GREY).

### 3.4.2 Allergen immunotherapy

- 1. Omalizumab (NICE TA278) for allergic asthma is **RED** NHSE commissioned.
- 2. Mepolizumab (NICE TA671) for eosinophilic asthma is **RED** NHSE commissioned.

### 3.4.3 Allergic emergencies

**Emerade** auto-injector 300 micrograms, 500 micrograms (see advice below) <u>PIL</u> **Jext** auto-injector 150 micrograms, 300 micrograms <u>PIL</u> **EpiPen** auto-injector 0.3mg, **EpiPen Jr** auto-injector 0.15mg <u>PIL</u> /<u>EpiPen Jnr PIL</u>

- See <u>MHRA June 2023</u>- Adrenaline auto-injectors (AAIs): new <u>guidance</u> and resources for safe useincluding an easy step-by-step guide on what to do in an emergency and updated advice on body positioning. A toolkit of resources is available for health and social care professionals to support the safe and effective use of AAIs. Use the materials to inform patients and caregivers what do if they suspect anaphylaxis and how to use adrenaline auto-injectors (AAIs)
- 2. <u>MHRA 2017</u> recommends two adrenaline auto-injectors should be prescribed which patients should carry at all times.
- 3. Adrenaline has a narrow therapeutic index. Primary care clinicians should prescribe in line with product licensing as summarised in the table below:-

	Weight range	Dose
Emerade	>30kg	300 micrograms
Jext	15-30kg	150 micrograms
	>30kg	300 micrograms
Epipen	Between 7.5 to 25kg	150 micrograms (Epipen Junior)
	>25kg	300 micrograms

- 4. The 500micrograms adrenaline dose (Emerade) should only be prescribed for self-administration on the advice of a specialist, for example where a repeated second dose of adrenaline (300microg) was necessary or in obese patients where a larger dose is necessary.
- 5. For educational material produced by manufacturer see links:- <u>Emerade</u>, <u>EpiPen</u>, <u>Jext</u> Other resources see links: <u>Emerade</u>, <u>Epipen</u>, <u>Jext</u> (key differences between devices)

## 3.7 Mucolytics

Carbocisteine	375mg caps, 750mg/10ml oral solution sachets
Acetylcysteine	600mg Sugar-free effervescent tablets

- 1. <u>NICE NG115</u> COPD in over 16s: diagnosis and management (Dec 2018):
  - Consider mucolytic drug therapy for people with a chronic cough productive of sputum.
  - Only continue mucolytic therapy if there is symptomatic improvement (for example, reduction in frequency of cough and sputum production).
  - Do not routinely use mucolytic drugs to prevent exacerbations in people with stable COPD.