



Indiana Medicaid Therapeutics Committee **Therapeutic Class Review Summary**

Therapeutic Class:

Oral inhaled corticosteroids

Overview:

Asthma is a chronic inflammatory disorder of the airways that contributes to airway hyperresponsiveness and airflow limitation. Airflow limitation is comprised of acute bronchoconstriction, airway edema, mucus plug formation, and airway wall remodeling. These features lead to bronchial obstruction. The immunohistopathologic features responsible for this inflammation process consist of denudation of airway epithelium, collagen deposition beneath basement membrane, edema, mast cell activation, and inflammatory cell infiltration by cells such as neutrophils, eosinophils, and lymphocytes.

Pharmacologic therapy helps prevent and control asthma symptoms, reduce the frequency and severity of exacerbations, and reverse airflow obstruction. Both acute and maintenance medications are used to achieve and maintain control of persistent asthma. Corticosteroids have consistently demonstrated effectiveness for long-term control of asthma by reducing symptom severity, improving peak expiratory flow, diminishing airway hyperresponsiveness, and potentially preventing airway wall remodeling.

The oral corticosteroid inhaler therapeutic class is comprised of several medications with numerous delivery systems. All agents within this class are indicated for the chronic treatment of asthma in adults while treatment ages for pediatric patients vary. The agents within this class are similar in terms of efficacy and safety. Inhaled corticosteroids are also available in combination with long-acting beta₂-adrenergic agonists. Advair Diskus[®]/ Advair[®] HFA (salmeterol/fluticasone), and Symbicort[®] (budesonide/formoterol) are the available combination products, and are discussed in a separate review. Asmanex[®] Twisthaler[®] (mometasone) and the recently approved agent, Alvesco[®] (ciclesonide) are indicated for the maintenance treatment of asthma as prophylactic therapy in patients 12 years of age and older. Asmanex is also indicated for the treatment of asthma patients who require oral corticosteroid therapy, where adding Asmanex therapy may reduce or eliminate the need for oral corticosteroids. Beclomethasone products Vanceril[®], Vanceril[®] DS, and Beclovent[®] are no longer available in the United States; however, QVAR[®] is available. Pulmicort Turbuhaler was recently phased out and replaced by Pulmicort Flexhaler, which comes in different strengths, and has a dose counter. Pulmicort Respules[®] are now approved and available in a generic formulation.

In April 2010, the The U.S. Food and Drug Administration announced, in accordance with longstanding U.S. obligations under the Montreal Protocol on Substances that Deplete the Ozone Layer, several metered-dose inhalers (MDI) used to treat asthma and chronic obstructive pulmonary disease (COPD) will be gradually removed from the U.S. marketplace. These inhalers contain ozone-depleting chlorofluorocarbons (CFCs), which are propellants that move medication out of the inhaler and into the lungs of patients. The ICS inhalers that will be phased out are Azmacort and Aerobid. Azmacort has already been discontinued by the manufacturer as of December 31, 2009 and is no longer available. Alternative medications that do not contain CFCs are available.

Generic Name	Brand Name	Pediatric Indication	Manufacturer
Beclomethasone Dipropionate HFA	QVAR [®]	≥ 5 Years of Age	IVAX
Budesonide	Pulmicort Respules [®] , Pulmicort Flexhaler [®]	≥ 1 Year of Age	AstraZeneca
Ciclesonide	Alvesco [®]	≥ 12 Years of Age	Nycomed
Flunisolide	AeroBID [®] , AeroBID [®] -M	≥ 6 Years of Age	Forest
Fluticasone	Flovent [®] Diskus, Flovent [®] HFA	≥ 4 Years of Age	Glaxo SmithKline
Mometasone Furoate	Asmanex [®] Twisthaler [®]	≥ 12 years of Age	Schering
Triamcinolone	Azmacort [®] (Discontinued)	≥ 6 Years of Age	Abbott

Summary:

The oral steroid inhalers include the following agents:

- beclomethasone dipropionate HFA (QVAR[®])
- ciclesonide (Alvesco[®])
- triamcinolone (Azmacort[®])
- flunisolide (AeroBID[®], AeroBID[®]-M)
- fluticasone (Flovent[®] Diskus, Flovent[®] HFA)
- budesonide (Pulmicort Flexhaler[®], Pulmicort Respules[®])
- mometasone furoate (Asmanex[®] Twisthaler[®])
- fluticasone/salmeterol (Advair[®] HFA)
- budesonide/formoterol (Symbicort[®])



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Azmacort[®] and AeroBID[®] are being discontinued, as they contain ozone-depleting CFCs. These agents decrease airway hyperresponsiveness by blocking late reaction to allergens, inhibit cytokine production, adhesion protein activation, and inflammatory cell migration and activation, reverse beta₂-receptor down-regulation, and inhibit microvascular leakage. The agents within this class are similar in terms of efficacy and safety; therefore, selection of a preferred agent should be based on other clinical considerations (dosing regimen, pediatric indication, etc.) and total cost impact to the health care system.