### IJPSR (2010), Vol. 1, Issue 12

## (Review Article)



INTERNATIONAL JOURNAL OF PHARMACEUTICAL SCIENCES AND RESEARCH



Received on 22 July, 2010; received in revised form 04 November, 2010; accepted 10 November, 2010

### ASTHMA AND HERBAL DRUGS

R. N. Kale<sup>\*1</sup> R. N. Patil<sup>1</sup> and R. Y. Patil<sup>2</sup>

SVPM's College of Pharmacy<sup>1</sup>, Malegaon BkII Tal- Baramati, Pune (Maharashtra), India PDEA's Shankarrao Ursal College of Pharmacy<sup>2</sup>, Kharadi, Pune (Maharashtra), India

#### Keywords:

Asthma, Medicinal plants, Bronchodilator, Mast cell stabilizer

#### Correspondence to Author:

#### R. N. Kale

SVPM's College of Pharmacy, Malegaon BkII Tal- Baramati, Pune (Maharashtra), India

Asthma is disease of the human respiratory system in which the airways constrict and become narrow, often in response to a "trigger" such as exposure to an allergen, cold air, exercise or emotional stress. Due to rapid industrialization and urbanization, asthma prevalence is predicted to increase more rapidly in the coming years. Despite the availability of a wide range of drugs for the treatment of asthma, the relief offered by them is mainly symptomatic and short lived. Moreover the side effects of these drugs are also quite disturbing. Medicinal plants have been known for millennia and are highly esteemed all over the world as a rich source of therapeutic agents for the prevention of diseases and ailments. The importance of herbal medicine in the treatment of asthma is indisputable. Four of the five classes of drugs currently used to treat asthma namely;  $\beta_2$ -agonists, anticholinergics, methylxanthines and cromones have origins in herbal treatments going back at least 5000 years. In the present article an attempt has been made to review antiasthmatic medicinal plants with their active chemical constituent and possible mechanism of action.

ABSTRACT

**INTRODUCTION:** The term "asthma" comes from the Greek meaning, "to breathe hard." The Global Initiative for Asthma was created to increase awareness of asthma among health professionals, public health authorities and the general public to improve prevention and management through a concerted worldwide effort. Bronchial Asthma according to the GINA guidelines final update November 2006 is clearly defined as: A chronic inflammatory disorder of the airways in which many cells and cellular elements play a role. The chronic inflammation is associated with airway hyper responsiveness that leads to recurrent episodes of wheezing, breathlessness, chest tightness and coughing, particularly at night or in the early morning. These episodes are usually associated with widespread, but variable, airflow obstruction within the lung that is often reversible either spontaneously or with treatment <sup>1</sup>.

Various research studies indicates that airway hyper responsiveness is important in the pathogenesis of asthma and the level of airway hyper responsiveness usually correlates with the clinical severity of asthma<sup>2</sup>. Based on the presence and absence of an underlying immune disorder asthma may be classified as a) extrinsic asthma in which asthmatic episode is initiated by type I hypersensitive reaction induced by an exposure to an extrinsic antigen and b) intrinsic asthma, in

which the triggering mechanisms are non immune and stimuli that have little or no effect in normal subjects can trigger bronchospasm<sup>3</sup>. Due to rapid and urbanization, industrialization asthma prevalence is predicted to increase more rapidly in the coming years. Although limited data is available on the asthma prevalence in India, according to the "Global Burden of Asthma Report", the increase is likely to be dramatic, particularly in India. A wide variation ranging from 4-19% is reported in the prevalence of asthma in school-going children from different parts of India. The prevalence of current-wheezing in children in Delhi is 16.7% and the cumulative prevalence is 20.8%<sup>4</sup>.

Pathophysiology of Asthma: Bronchial asthma is characterized pathologically by an infiltration of eosinophils into the airway submucosa. Eosinophil activation results in the secretion of an array of highly charged cytotoxic cationic proteins such as major basic protein, and is believed to play a central role in the etiology of this disease by inducing damage to the airway epithelium  $^{5}$ . The pathophysiology of asthma involves the development of acute and chronic inflammation in airway narrowing by producing increased vascular permeability, edema, and airway smooth muscle contraction<sup>6</sup>.



IG.1: PATHOPHYSIOLOGY OF ASTHMA

Available online on www.ijpsr.com

The gross pathology of asthmatic airways displays lung hyperinflation, smooth muscle hypertrophy, lamina reticularis thickening, mucosal edema, epithelial cell sloughing, cilia cell disruption, and mucus gland hypersecretion<sup>7</sup>. It is observed that those patients who have died due to asthma are due to considerable increase in the thickness of the airway wall throughout the bronchial tree, partly as a result of smooth muscle hypertrophy<sup>8</sup>.

**Treatment:** The following category of drugs can be used alone or in combination for the treatment of asthma <sup>10</sup>.

# 1. Bronchodilators

- Beta 2 Adrenergic agonists
- Muscarnic antagonists
- Methyl xanthines

# 2. Anti inflammatory agents

- Glucocorticoids
- Mast cell degranulation blockers (Mast cell stabilizers)

# 3. Newer drugs

- Leukotriene antagonists
- Anti I<sub>g</sub> E antibodies
- Allergy vaccination

Herbal Drugs used for Asthma: Though the large numbers of drugs are available for the treatment of asthma, the relief offered by them is mainly symptomatic and short lived. Moreover the side

### TABLE 1: BRIEF REVIEW OF ANTIASTHMATIC PLANTS

effects of these drugs are also quite disturbing. Recently there has been a shift in universal trend from synthetic to herbal medicine, which we can say 'Return to Nature'. Medicinal plants have been known for millennia and are highly esteemed all over the world as a rich source of therapeutic agents for the prevention of diseases and ailments <sup>11</sup>. A large number of medicinal plants have been used traditionally for the treatment of asthma and have been scientifically proven to have antiasthmatic properties.

Important medicinal plants having antiasthmatic potential are Achyranthes aspera, <sup>12</sup> Allium cepa, <sup>12, 13</sup> Adhatoda vasica, <sup>14, 15</sup> Albizzia lebbeck, <sup>16, 17</sup> Achillea mellifolium,<sup>18</sup> Asystasia gangetica,<sup>19</sup> Acorus calamus, Ammi visnaga, Boswellia serrata,<sup>20</sup> Balanites roxburghii,<sup>21</sup> Cedrus deodara,<sup>22</sup> Curculigo orchioides,<sup>23</sup> Clerodendron phlomidis,<sup>24</sup> Curcuma longa,<sup>25</sup> Cassia sophera,<sup>26</sup> minima,<sup>27</sup> Ephedra globules,<sup>29</sup> Aeqle gerardiana,<sup>28</sup> Centipeda marmelos.<sup>30</sup> Eucalyptus Hedychium spicatum,<sup>31,32</sup> Glycyrrhiza glabra,<sup>33,34</sup> Inula racemosa,<sup>35</sup> , Moringa oleifera,<sup>36</sup> Myrica sapida,<sup>37</sup> Nigella sativa, Ocimum sanctum, Picorrhiza kurroa,<sup>38</sup> Lipidum sativum,<sup>39</sup> Passiflora incarnata,<sup>40</sup> xanhocarpum,<sup>41,</sup> Solanum Terminalia belerica,<sup>43</sup> Tinospora cordifolia,<sup>44</sup> Tamarandus indica <sup>45</sup>.

Following **table 1** gives a brief review of the antiasthmatic plant with their chemical constituent and probable Mechanism of action.

| PLANTS             | FAMILY        | PART USED    | CHEMICAL CONSTITUENTS | MECHANISM OF ACTION  |  |  |
|--------------------|---------------|--------------|-----------------------|--|--|--|
| Achyranthes aspera | Amaranthaceae | Fruit        | Saponin C Saponin D   | Mast cell stabilizer   |  |  |
| Allium cepa        | Liliaceae     | Bulb         | Quercetin             | <ol> <li>Mast cell stabilizer</li> <li>Lipoxygenase inhibitor</li> <li>PAF inhibitor</li> <li>COX inhibitor</li> </ol> |  |  |
| Adhatoda vasica    | Acanthaceae   | Leaves, root | Alkaloids             | <ol> <li>Bronchodilator</li> <li>Anti-anaphylactic</li> </ol>  |  |  |

Available online on www.ijpsr.com

# International Journal of Pharmaceutical Sciences and Research

ISSN: 0975-8232

| Albizzia lebbeck       | Leguminosae             | Bark           | Alkaloids,                         | 1. Bronchodilator                            |
|------------------------|-------------------------|----------------|------------------------------------|--|
|                        | -                       |                | tannins, navonoids,                | 2. Mast cell stabilizer                      |
| Achillea mellifolium   | Asteraceae (compositae) | Flower         | Alkaloids                          | acetylcholine and 5-HT                       |
| Asustasia gangotica    | Aconthocopo             | Loovos         | Triterpenoids, saponins, Steroidal | 1. Bronchodilator                            |
| Asystusia gungetica    | Acanthaceae             | Leaves         | aglycone                           | 2. Anti-inflammatory                         |
| Acorus                 |                         |                |                                    | Inhibits action of histamine.                |
| calamus                | Araceae                 | Rhizome        | Asarone                            | acetylcholine and 5-HT                       |
| Ammi visnaga           | Umbelliferae            | Seeds          | Khellin                            | Bronchodilator                               |
| Boswellia serrata      | Burseraceae             | Root           | Boswellin,                         | Inhibits leukotriene                         |
|                        |                         |                | Boswellic acid                     | biosynthesis                                 |
| Balanites roxburghii   | Simarubaceae            | Stem bark      | Alkaloids                          | 1. Bronchodilator                            |
| -                      |                         |                |                                    | 2. Mast cell stabilizer                      |
| Cedrus deodara         | Pinaceae                | Wood           | Himacholol                         | Mast cell stabilizer                         |
| Curculigo orchioides   | amarylliaceae           | Rhizomes       | Triterpenoids sapogenins and       | 1. Antihistaminic                            |
|                        |                         |                | saponin grycosides                 | 2. Anti-initatimatory                        |
| Clerodendron phlomidis | Verbenaceae             | Leaves         | Flavonoids, terpenoids, steroids   | 1. Antihistaminic<br>2. Mast cell stabilizer |
|                        |                         |                |                                    | 2. What cell stabilizer                      |
| Curcuma longa          | Zingiberaceae           | Rhizome        | Curcuminoids                       | Inhibits histamine release                   |
|                        |                         |                |                                    | 1. Bronchodilator                            |
| Cassia sophera         | Caesalpiniaceae         | Leaves         | Flavonoids, glycosides             | 2. Antihistaminic                            |
| cussia sopricia        |                         |                |                                    | 3. Antiallergic                              |
|                        |                         |                | Pseudoguainolide, sesquiternene    | 4.anti-initatinitatory                       |
| Centipeda minima       | Compositae              | Whole plant    | lactone, flavonoids                | Antiallergic                                 |
| Ephedra gerardiana     | Ephedraceae             | Stem           | Ephedrine                          | Bronchodilator                               |
| Eucalyptus globules    | Myrtaceae               | Leaves         | Volatile oil                       | Anti-inflammatory                            |
| Aegle marmelos         | Rutaceae                | Leaves         | Alkaloid-aegeline                  | Antihistaminic                               |
| Hedychium spicatum     | Zingiberaceae           | Rhizome        | Sitosterol, Volatile oil           | Anti inflammatory                            |
| Glycyrrhiza glabra     | Leguminosae             | Root           | Glycyrrhizinic acid                | 1.Antihistaminic                             |
| Inula racomoca         | Astorosooo              | Deete          | Inulin, sesquiterpene lactone-     | Antihistominic                               |
| mulu rucemosu          | Asteraceae              | ROOIS          | alantolactone                      | Antinistaminic                               |
| Moringa olaifara       | Morangacoao             | Sood           | Tannins, steroids, triterpenoids,  | Antihistominic                               |
| woringa ölelfera       | Wordigaceae             | Seeu           | flavonoids, alkaloids, saponins    | Antinistaninin                               |
| Myrica sapida          | Myricaceae              | Bark           | Glycosides                         | Mast cell stabilizer                         |
| Nigella sativa         | Ranunculaceae           | Seed           | Volatile oil, fatty acid           | Bronchodilator                               |
| Ocimum sanctum         | Labiateae               | Leaves         | Ursolic acid                       | Mast cell stabilizer                         |
| Picorrhiza kurroa      | Scrophulareaceae        | Roots          | Picorrhizin                        | Antihistaminic                               |
| Lipidum sativum        | Cruciferae              | Seeds          | Alkaloids, Flavonoids              | Bronchodilator                               |
| Passiflora incarnata   | Passifloraceae          | Leaves         | Benzoflavone                       | Bronchodilator                               |
| Colomation             | Colone                  | <b>Flauren</b> | Phyto-sterol, alkaloids,           | 1. Antihistaminic                            |
| Solanum xanhocarpum    | Solanaceae              | Flowers        | flavonoooids, Steroids             | 2. Mast cell stabilizer                      |

Available online on www.ijpsr.com

| Terminalia belerica  | Combrataceae    | fruits | Beta sitosterol, Gallic acid, ellagic<br>acid, glycoside | Mast cell stabilizer  |
|----------------------|-----------------|--------|--|---|
| Tinospora cordifolia | Mensipermaceae  | Stem   | Alkaloids  | <ol> <li>Antihistaminic</li> <li>Mast cell stabilizer</li> </ol>                              |
| Tamarindus indica    | Caesalpiniaceae | leaves | Flavone, Glycosides                                      | <ol> <li>1.Brochodialator</li> <li>2. Antihistaminic</li> <li>3. Anti-inflammatory</li> </ol> |

**CONCLUSION:** Many synthetic drugs are used to treat asthma, but they are not completely safe for long term use. Nature has bestowed our country with an enormous wealth of medicinal plants; therefore India has often been referred to as the Medicinal Garden of the world. Scientifically explored exhaustive reports published in Indian and international journals suggest the importance of herbal medicine in the treatment of asthma is indisputable.

#### **REFERENCES:**

- Tarek S. Towards a deep understanding of bronchial asthma. Egyptian Journal of Bronchology, 2007 Dec; 1(1): 120-124
- Busse WW, Calhoun WJ, Sedgwick JD. Mechanisms of airway inflammation in asthma. Am. Rev. Respir. Dis. 147:S20-S24
- 3. Doshi U, Salat P, Parikh V. Cytokine modulators in Asthma: clinical perspectives. Indian J pharmacol 2002; 34: 16-25
- Goswami DV, Sonawane LL, Nirmal SA, Patil MJ. Evaluation of antiasthmatic activity of *tectona grandis* linn. Bark. Int J Pharm. Sci. and Res.2010;1(1):10-16
- Christopher EB, Peter B, Fiona AS, Stephen TH, Andrew J W, Ian DP. Mast-cell infiltration of airway smooth muscle in asthma. *New Eng J Med* 2002; 33; 1699-1705.
- 6. Gregory GK, Peter DP, Chun YS. The mechanics of exaggerated airway narrowing in asthma the role of smooth muscle. *Respiration Physiology 1999*; 118:1-13.
- 7. Sundeep SS, Mamudipudi TK, Anthony PS, Stephen TH. The anti-inflammatory effects of leukotriene-modifying drugs and their use in asthma. *Chest* 2001; 119:1533-1546
- Wardlaw AJ, Brightling CE, Green R, Woltmann G, Bradding P, Pavord ID. New insights into the relationship between airway inflammation and asthma. *Clinical Science* 2002; 103: 201–211
- 9. from-http://www.diabetesjain.com/asthma.gif
- 10. Rajani GP, Patil UA, Deb J. Anti asthmatic an overview Indian Drugs 2006; 43(10):781-89
- 11. Sharma A, Shanker C, Tyagi L, Singh M, Rao CV. Herbal medicine for market potential in India: An Overview. Academic Journal of Plant Sciences. 2008; 1(2): 26-36.

- Rastogi RP, Mehrotra BN, Compendium of indian medicinal plants, 1st Edition, Vol 3, CDRI Lucknow and PID New Delhi, 1980-1984; 10, 224, 294, 376
- Chaudhari RD (Ed.). Pharmacological classification of medicinal herbs, Herbal Drug industry, Eastern Publishers, New Delhi, 1st Edition, 2004; 61.
- Paliwa JK, Dwiwedi AK, Singh S. Pharmacokinetics and insitu absorption studies of a new antiallergic compound 73/602 in rats, Int. J. Pharm. 2000;197(1-2): 213-220
- 15. Evans WC. Trease and Evans Pharmacognosy, 15th edn, WB Saunders Company Ltd., London, 2003; 299, 471, 485
- 16. Tripathi RM, Das PK. Studies on antiasthmatic and anti anaphylactic activity of *Albizzia lebbeck* Ind. J. Pharmacol 1977, 9(3):189-194
- 17. Tripathi RM, Sen PC, Das PK Studies on the mechanismof action of *Albizzia lebbeck*, an indigenous drug used in the treatment of atopic allergy, J Ethnopharmacol.1979,1, 1385-1396.
- Raju D, Chitra V, Hari Das K, Silambu Janiki P, Shankari M. Evaluation of antiasthmatic activity of aqueous extract of *Achillea mellifolium linn* flowers. Arc. Apl. Sci. Res., 2009 1(2): 287-293
- 19. Akah PA, Ezike AC, Nwafr SV, Okoli CO, Enwerem NM. Evaluation of the antiasthmatic property of *Asystasia* gangetica leaf extracts. J Ethnopharmacol. 2003 (89):25-36.
- Gupta I, Gupta V, Parihar A, Gupta S, Ludtke R, Safeyhi H, Ammon HP. Effects of *Boswellia serrata* gum resin in patients with bronchial asthma: results of a double-blind, placebo-controlled 6 week clinical study. Eur. J. Med. Res. 1998; (11): 511-514
- 21. Singh V, Tripathi P, Patel JR, Kori ML, Dixit VK, Preliminary phytochemical and antiasthmatic studies on stem bark of *Balanites roxburghii* planch. International Journal of Pharmaceutical and Clinical Research 2009,1(1):40-42
- Shinde UA, Phadke AS, Kulkarni KR, Nair AM, Mungantiwar AA, Dixit VJ, Saraf MN Mast cell stabilizing and lipoxygenase inhibiting activity of *Cedrus deodara*(Roxb.) wood oil. Indian J.Exp.Biol. 1999; 37(3): 258-261.
- Pandit P, Singh A, Bafna AR, Kadam PV, Patil MJ. Evaluation of antiasthmatic activity of Curculigo orchioides Gaertn. Rhizomes. Ind J Pharm Sci. 2008 July-August; 440-444
- 24. Vadnere GP, Somani RS, Singhai AK. Studies on antiasthmatic activity of aqueous extract of *Clerodendron phlomidis*. Pharmacologyonline. 2007(1):487-494
- 25. Ammon HP, Wahl MA. Pharmacology of curcuma longa. Planta Medica 1991,57(1), 1-7

Available online on www.ijpsr.com

- Nagore DH, Ghosh VK, Patil MJ. Evaluation of antiasthmatic activity of *Cassia sophera* Linn. Pharmacognosy magazine. 2009; 5(19):109-118
- 27. Wu JB, Chun YT, Ebizuka Y, Sankawa V. Biologically active constituents of *Centipeda minima*: isolation of a new phenolin ester and the antiallergic activity of sesquiterpene lactones. Chem. Pharm. Bull.1985,33: 4091-4094
- Rastogi RP, Mehrotra BN. Compendium of Indian Medicinal Plants, 1st Edition, Vol 4, CDRI Lucknow and PID New Delhi, 1985-1989; 154, 246, 288, 315, 348, 360.
- 29. Focho DA, Nkeng EAP, Fonge BA, Fongod AN, Muh CN, Ndam TW, Afegenui A. Diversity of plants used to treat respiratory diseases in Tubah, northwest region, Cameroon. African J. of Pharm. Pharmacol. 2009 Nov3(11): 573-580
- Arul V, Miyazaki A, Dhananjayan R. Mechanism of the contractile effect of the alcoholic extract of *Aegle marmelos* corr. on isolated guinea pig ileum and tracheal chain. Phytomedicine. 11(7): 679-683
- 31. Chaturvedi GN, Sharma BD. Ethnobotnical survey of the plants used to treat asthma in Andhra Pradesh. Journal of research in Indian medicine. 1975;10(2):6
- Shekhar AV, Gandhi DN, Mohan Rao N, Rawal UD. An experimental and clinical evaluation of Antiasthmatic potentialities of Devadaru compound (Dc) Indian J Physiol Pharmacol. 2003 Jan.47(1): 101-107
- RP Rastogi, BN Mehrotra. Compendium of Indian Medicinal Plants, 1st Edition, Vol 4, CDRI Lucknow and PID New Delhi, 1985-1989; 154, 246, 288, 315, 348, 360.
- 34. WC Evans. Trease and Evans Pharmacognosy, 15th edn, WB Saunders Company Ltd., London, 2003; 299, 471, 485.
- Srivastava S, Gupta PP, Prasad R, Dixit KS, Palit G, Ali B, Misra G, Saxsena RC. Evaluation of antiallergic activity (type I hypersensitivity) of *Innula racemosa* in rats. Indian J. Physiol. Pharmacol. 1999;29,89-95

- Agrawal B, Mehta A .Antiasthmatic activity of *Moringa* olifera Lam: A clinical study Indian J Pharmacol. 2009; 40(1): 28-31
- Patel KG, Bhalodia PN, Patel AD, Patel KV, Gandhi TR. Evaluation of bronchodilator anti anaphylactic activity of *Myrica sapida*. Iranian Biomedical Journal .2008;12 (3):191-196
- Doshi VB, Shetge VM, Mahashur AA, Kamat SR, *Picrorrhiza kurroa* in bronchial asthma. J. Postgrad. Med.1983;29:89-95
- Mali RG, Mahajan SG, Mehta AA. Studies on bronchodilatory effect of *Lepidium sativum* against allergen induced bronchospasm in guinea pigs. Pharmacognosy magazine. 2008; 4(15):189-192
- 40. Dhawan K, Kumar S, Sharma A. antiasthmatic activity of the Methanol extract of leaves of *Passiflora incarnata*.Phytother.Res.2003;17:821-22
- 41. Vadnere GP, Gaud RS, Singhai AK. Evaluation of antiasthmatic property of *Solanum xanthocarpum* flowers extracts. Pharmacologyonline. 2008;1: 513-522
- Govindan S, Vishwanathan S, Vijaysekaran V, Alagappan R, A pilot study on the clinical efficacy of *Solanum xanthocarpum* and *solanum trilobatum* in bronchial asthma. J Ethnopharmacol. 1999;66: 205-210
- 43. Vyas BA, Vyas RB. Effect of ethanolic extracts of *Allium* sativum and *Terminalia belerica* on clonidine induced mast cell degranulation and clonidine and haloperidol induced catalepsy in mice. Int J Pharm Res 2009; 1(1) : 41-44
- 44. Nayampalli SS, Desai NK, Ainapure SS.Antiallergic properties of *Tinspora cordifolia* in animal models. Indian J. Pharmacol. 1986;18:250-252
- 45. Tayade PM, Ghaisas MM, Jagtap SA, Dongre SH. Antiasthmatic activity of methanolic extract of leaves of *Tamarandus indica* linn. Journal of pharmacy research.2009, 2(5):944-947

\*\*\*\*\*\*