Prospects of botanical dewormers with especial reference to Kashmir

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DOI: https://doi.org/10.22271/veterinary.2021.v6.i4a.372

Abstract
The increasing prevalence of anthelmintic resistant strains of helminth parasites, drug residues in animal products and high cost of marketed anthelmintics has enhanced an attention in studying medicinal plants as an alternative source of anthelmintics. Alkaloids, alpha-pinene, artemisin, benzyl isothiocyanate, Cucurbitacin, dithymoquinone, genistein, papain, phenolic compound, tannin and thymoquinone are some active constituents found in different plants and act against the helminth parasites of animals and human beings. Mostly the tannin-rich forages offer an alternative to anthelmintic chemicals in controlling gastrointestinal nematodes. The traditional use of different plants showing the anthelmintic activities against various endoparasites of animals in different regions of country with special reference to the Kashmir valley are reviewed in this communication.

Keywords: Carcass characteristics, weaner pigs, meat quality, palm kernel cake

Introduction
India possesses largest livestock population in the world which plays an important role in Indian economy. The livestock sector has a major significance in the rural economy of Jammu and Kashmir contributing 5.04% to the state GDP [1]. Despite the great contribution of livestock sector to the economy and livelihood of the people of Kashmir valley it is further aggravated by various bacterial, viral and parasitic diseases. Parasitic diseases are considered to be the most prevalent and important health problems in grazing ruminants inflicting losses through mortality, feed efficiency ratio and by way of costs incurred upon their treatment and control [2]. Amongst the helminth parasites, gastrointestinal (GI) parasite infected animals have lower growth rates, reduced reproductive performance, and have higher rates of illness and death. Therefore anti-parasitic drugs are used to control the parasitic diseases. An ideal anthelmintic drug should have broad spectrum of action, high cure rate with single dose, free from toxicity to the host and also cost effective. But, no synthetic anthelmintic agent meets all requirements. Ethno-veterinary practices can be used as sustainable development through appreciating and building upon local knowledge of veterinary and husbandry practices. In this way, herbal medicine may very well become a pioneer in turning local knowledge into global knowledge, through the recognition of local knowledge as an indispensable source of sustainable development for both people and animals all over the world. Traditional veterinary medicine has been practised as early as 1800 B.C. at the time of king Hammurabi of Babylon [3]. Therefore, switch to ethno-veterinary drugs is based on the cultural heritage of an area that involves interaction between plants and the people including the traditional use of medicinal plants and their management is referred as ethnobotany [4]. A plant with therapeutic properties naturally synthesise and accumulates some secondary metabolites like alkaloids, volatile oils, vitamins and minerals in different parts such as leaves, fruits, seeds and rhizomes [5,6] and play a significant role in providing health care and improving economy of the farmers [7]. Satyavati et al. [8] reported the plant kingdom is known to provide a rich source of botanical anthelmintics, anti-bacterial and insecticides. A number of medicinal plants have been used to treat parasitic infections in man and animals. There are many plants which have been reported in the literature for their anthelmintic importance. A number of studies have been carried out showing that certain plant species have the nutritional value but also show promising results in reducing the parasitic infection in sheep [9, 10].
Traditional use of plants against parasitic infections in human beings as well as livestock is common practice in Kashmir valley. A number of compounds have been identified which show promising anti-parasitic activity. Some of them are discussed below.

**Alkaloids**

Alkaloids present in different plants used against different helminth parasites may act on central nervous system and cause paralysis of the helminthes. Steroidal alkaloid and oligoglycosides are present in alkaloids which may suppress the transfer of sucrose from the stomach to the small intestine. Alkaloids also act as an antioxidant, capable of reducing the nitrate generation which can interfere in local homeostasis that is essential for the development of helminths [11].

**Phenolic compound**

Phenolic compounds present in different parts of different plants when administered against endoparasites interfere with the energy generation mechanism by uncoupling the oxidative phosphorylation and also interfere with the glycoproteins of the cell surface of the parasites and cause their death [12].

**Tannins**

Various pasture tanniferous plants have been investigated for potential effect against either incoming parasite larvae and/or already established worms [13]. Tannins interfere with energy generation of worms by uncoupling oxidative phosphorylation or they binds to the free proteins of the gastrointestinal tract of the host animal or glycoproteins on the cuticles of the worms and lead to death [14]. The beneficial effects of tanniferous plants against internal parasites could be due to one, or a combination, of the following factors:

1. Tanniferous plants increase the supply and absorption of digestible protein by animals. This is achieved by tannins forming non-biodegradable complexes with protein in the rumen, which dissociate at low pH in the abomasum to release more protein for metabolism in the small intestine of ruminants. This indirectly improves host resistance and resilience to nematode parasite infections [15].
2. Tannins have a direct anthelmintic effect on resident worm populations in animals [15].
3. Tannins and/or their metabolites in dung have a direct effect on the viability of the free-living stages (development of eggs to infective larval stages).

These plants can be a promising future for the control of worms which had previously shown resistance to synthetic drugs. These plants can be used to feed animals not only as source of nutrition but also as a source of certain antiphrastic substances. It is well known that tannins can produce this effect; however, similar results can be obtained with a variety of other substances. Tannins are usually divided into hydrolysable and condensed tannins (CT), although there are other types.

Condensed tannins, the most frequent type of tannins found in leguminous plants and trees are relatively stable in the digestive tract of the animals and they rarely have toxic effects and can therefore, used to control parasite populations. CT are a diverse group of plant secondary metabolites usually found in leaves, roots, nuts and fruits from a wide-range of different plant sources in both tropical and temperate areas [16]. Tannins used in different concentrations (5-50% of DM) in animal diets can reduce the parasitic population. CTs have antimicrobial properties also [17]. Besides this, they have effect on *Candida albicans* [18], *Trypanosoma brucei* [19] and *Leishmania donovani* [20].

In addition, a number of studies conducted with nematode parasites of ruminants have demonstrated direct anthelmintic effects of extracts from tannin-containing plants in *in vitro* assays, with *in vivo* verification of these results being reported in some studies [21]. *Haemonchus contortus* infections in livestock have been found to be influenced by tannin structure [22] obtained from the diverse nature of CT in different plant sources. It was found that extracts of tannin-rich plants interfere with the first phase of host invasion, i.e., the exsheathment of infective third stage larvae [22].

Tannins have both direct and indirect effects over the gastrointestinal parasites in ruminants [19].

i. Direct effects: caused by the tannin-parasite interaction, affecting the parasite’s physiology.

ii. Tannins extracted from several forages reduce the total nematode burden, the migration, the viability of the larvae and the egg per gram (EPG) count.

iii. They can also interfere with the hatching of the eggs and the development of infective larval stages.

iv. CT of *Quebracho* had a direct anthelmintic effect against *Trichostrongylus colubriformis* infection in sheep.

v. The administration of quebracho extract reduces the EPG count, the adult burden and the fecundity of *T. colubriformis*.

vi. Indirectly, tannins improve the protein nutrition by binding themselves to the proteins of the plants in the rumen and preventing the microbial degradation, thus increasing the offer of amino acids to the duodenum.

vii. Recent experiments suggest that the addition of *quebracho* tannins to diets with low protein levels can improve the diet quality and reduce the consequences of parasitism.

**Among the most common medicinal plants which have anthelmintic effect are**

- *Allium sativum*, (garlic)
- *Nigella sativa*, (Kalonji, black cumin/ black caraway)
- *Artemisia spp.*, (thethwan)
- *Balanites aegyptiaca*, (desert date or Azores)
- *Acacia spp.*, (babul, kikar or Indian gum Arabic tree)
- *Cucurbita pepo* (pumpkin seeds),
- *Commiphora molmol* (Myrrh),
- *Calendula micrantha officinalis*, (marigold)
- *Peganum harmala* (Harmal/ Isband) and
- *Curcuma longa* (Tumeric) [23, 24]
- *Achillea millefolium* (Pahel gassa) Crude aqueous and ethanolic extracts have anthelmintic activity in sheep infected with mixed species of gastrointestinal nematodes.
Table 1: Medicinal plants used against helminth parasites of animals

<table>
<thead>
<tr>
<th>Plant</th>
<th>Part used</th>
<th>Active constituent</th>
<th>Indication cited or activity</th>
<th>Animal species</th>
<th>Reference</th>
</tr>
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<tr>
<td>Acacia albida</td>
<td>Seeds</td>
<td></td>
<td>-</td>
<td>Sheep, goat</td>
<td>Nwude and Ibrahim [55]</td>
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<tr>
<td>Langium lamarckii</td>
<td>Root-bark</td>
<td>Arscids</td>
<td>-</td>
<td>Poultry</td>
<td>Dubey and Gupta [32]</td>
</tr>
<tr>
<td>Albizia anthelmintica</td>
<td>Bark Root</td>
<td></td>
<td>-</td>
<td>Minja [27], ITDG and IIRR [28]</td>
<td></td>
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<tr>
<td>Albizia coriacea</td>
<td>Bark Bulb</td>
<td>Fascioloisis, lungworms</td>
<td>-</td>
<td>Cattle, goat, sheep, camel</td>
<td></td>
</tr>
<tr>
<td>Allium sativum</td>
<td>Leaves</td>
<td>Anthraquinone</td>
<td>Nippostrongylus brasiliensis</td>
<td>Rat</td>
<td>Ibrahim et al. [59]</td>
</tr>
<tr>
<td>Aloe barteri</td>
<td>Fruit</td>
<td></td>
<td>-</td>
<td>Chicken</td>
<td>Fernandez [36]</td>
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<td>Ananas comosus</td>
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<td>Annona cherimolia</td>
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<td>A. senegalensis</td>
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<td>Anogeissus leucarpus</td>
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<td>Areca catechu</td>
<td>Nut</td>
<td>Arecolin, other alkaloids</td>
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<td>Cattle, dog, goat</td>
<td>Roepeke [32]</td>
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<tr>
<td>Artemisia maritima</td>
<td>Santonin, Artemisin</td>
<td></td>
<td>Neocularis vitulorum</td>
<td>Buffalo calves</td>
<td>Akhtar et al. [37], Farnsworth et al. [34], Sherif et al. [37]</td>
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<tr>
<td>Bixa orellana</td>
<td>Seeds</td>
<td></td>
<td>Aescaria galli, Aescaria suum</td>
<td>Chicken, Pig</td>
<td>Fernandez [30]</td>
</tr>
<tr>
<td>Boswellia daltzii</td>
<td>Bark</td>
<td></td>
<td>Antihelminthic</td>
<td>Sheep, goat</td>
<td>Nwude and Ibrahim [32]</td>
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<tr>
<td>Butea frondosa</td>
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<td></td>
<td>Oxyurids</td>
<td>Mice</td>
<td>Mehta and Parashar [38], Lal et al. [37]</td>
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<td>Caesalpina cristia</td>
<td>Seeds</td>
<td></td>
<td>Toxoacra vitulorum, Aescaria galli,</td>
<td>Buffalo calves, chicken</td>
<td>Akhtar et al. [58]</td>
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<td>Collinida portoricensis</td>
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<td>Toxoacra canis</td>
<td>Dog</td>
<td>Adewumi and Akuba [38]</td>
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<tr>
<td>Carica papaya</td>
<td>Latex from fruit</td>
<td></td>
<td>Aescaria galli, Aescaria suum,</td>
<td>Chicken, Pig, Mice</td>
<td>Mursof and He [49], Satrija et al. [42]; Satrija et al. [42]</td>
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<td>Cassida eduvis</td>
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<td></td>
<td>Heligmososomatus polyergus</td>
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<td>ITDG and IIRR [36]</td>
</tr>
<tr>
<td>Cassia alata</td>
<td>Seeds</td>
<td></td>
<td>Aescaria galli</td>
<td>Chicken</td>
<td>Fernandez [32]</td>
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<td>Chrysanthemum spp.</td>
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<td></td>
<td>Aescaria lineata</td>
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<td>Rebrasser [47]</td>
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<tr>
<td>Diospyros mollis</td>
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<td></td>
<td>Necator americas, Nematodirus dubius,</td>
<td>Mice, Golden hamster</td>
<td>Sen et al. [44]</td>
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<tr>
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<td>Aescaria galli</td>
<td>Chicken</td>
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<td></td>
<td>Haemonchus contortus</td>
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<td>Fernandez [32]</td>
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<tr>
<td>Peganum harmala</td>
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<td></td>
<td>Tetra-hydro- harmun</td>
<td>Goat</td>
<td>Akhtar and Ahmad [41]</td>
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<tr>
<td>Quisqualis indica</td>
<td>Stem</td>
<td></td>
<td>Mixed gastro-intestinal infection</td>
<td>Goat, Pig</td>
<td>Fernandez [32], Farnsworth et al. [34]</td>
</tr>
<tr>
<td>Rupanea melanoploeos</td>
<td>Seeds</td>
<td></td>
<td>Aescaria suum, Aescaria galli</td>
<td>Chicken</td>
<td>Fernandez [32]</td>
</tr>
<tr>
<td>Rhus velaris</td>
<td>Root</td>
<td></td>
<td>Roundworms</td>
<td></td>
<td>ITDG and IIRR [36]</td>
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<tr>
<td>Tamarindus indica</td>
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<td></td>
<td>Roundworms</td>
<td></td>
<td>ITDG and IIRR [36]</td>
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<tr>
<td>Terminalia avicennoides</td>
<td>Leaf, root</td>
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<td>Anthraquinone</td>
<td>Nippostrongylus braziiliensis</td>
<td>Rat, Ibrahim [59]</td>
</tr>
<tr>
<td>Tinospora rhuphi</td>
<td>Stem</td>
<td></td>
<td>Haemonchus contortus</td>
<td>Goat</td>
<td>Fernandez [32]</td>
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<tr>
<td>Tribulus terrestris</td>
<td>Plant</td>
<td></td>
<td>Aescaria galli</td>
<td>Poultry</td>
<td>Chakraborty et al. [42]</td>
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<td>Uvaria hookeri</td>
<td>Root-bark</td>
<td></td>
<td>Acetogenins</td>
<td>Haemonchus contortus</td>
<td>Padmaja et al. [44]</td>
</tr>
<tr>
<td>Vernonia anthelmintica</td>
<td></td>
<td></td>
<td>Oxyurids</td>
<td>Mice</td>
<td>Mehta and Parashar [38]</td>
</tr>
</tbody>
</table>

Commiphora molmol (Myrrh)

Myrrh, an oleo-gum resin obtained from the stem of the plant species Commiphora (C. molmol) used as safe effective treatment for dicroellosis in sheep (Al-Mathal and Fouad, 2004) [48]. Myrrh used against schistosomosis in human beings showed that the drug was well tolerated, and side effects were mild and transient [49, 50]. Consequently, Myrrh extract of the medicinal plant, C. molmol proved to be safe and very effective in sheep monieziosis caused by Moniezia expansa [41].

Ocimum sanctum Linn. (Babri seeds)

It’s also known as sacred basil. The essential oil contains eugenol, B-caryophyllene and other monoterpenes in the plants which showed potent in-vitro anthelmintic activity [52, 53].

Melia azedarach (commonly known as the chinaberry tree)

Ethanolic extract of chinaberry, the plant which is native to India, China and Paris were found to act against tapeworms. Szewezuk et al. [54] found more efficacy of ethanolic extract of the plants as compare to piperaazine phosphate.

Artemisia annua (Tethwan)

It is also known as sweet wormwood, sweet anise, sweet sagwort which contains artemisin. Cumming et al. [55] reported that artemisinin damaged biological macromolecules of the parasites by causing oxidative stress in the cells; it is done by cleavage of endoperoxide bridges by iron producing free radicals (hypervalent iron-oxo species, epoxides, aldehydes and dicarbonyl compounds). Hence, the artemisin, the active compound of this tree may be used as plant dewormer against some internal parasites of animals [56]. Artemether, a methyl ether derivative has proven efficacy against Schistosoma haematobium, S. Japonicum and S. mansoni [57].

Carica papaya Linn.

Papain and Benzyisothiocyanate is the active principles of papaya which effect against helminta parasites of animals and human beings. Among these, benzy isothiocyanate which inhibits energy metabolism and affects motor activity of the parasites, hence acts as an anthelmintic agent [58]. Ethanolic extract of papaya were found effective against Paramphistomum cervi, H. contortus, Ascaris lumbricoides and Ascaridia galli [57, 59]. While papain, also known as papaya proteinase I, is a cysteine proteinase enzyme present in leaves, fruits, and seed of papaya (C. papaya) and mountain papaya (Vasconcellea pubescens). Papain consists of a single polypeptide chain with 3 disulphide bridges and a sulphydrol group necessary for activity of the enzyme which is responsible for digestion of nematodes cuticle.

Nigella sativa (kala jira)

~ 60 ~
Along with anthelmintic property it has digestive, carminative, diuretic and astringent attributes. The active ingredients of kala jira are thymoquinone, dithymoquinone and alpha-pinene as ingredients. N. sativa and its essential oil have anthelmintic action against trematodes, nematodes and cestodes [60, 61]. Black seeds of N. sativa induce oxidative stress against mature worms which was revealed by declining activities of antioxidant enzymes, glutathione peroxidase, glutathione reductase and superoxide dismutase as well as hexokinase and glucose-6-phosphate dehydrogenase, the glucose metabolism enzymes [62]. This plays a key role in the control of the overgrowth of helminthes [63]. N. sativa oil exhibits an additive effect in the treatment of schistosomiasis when it is given with praziquantel.

**Flemingia vestita** (Fabaceae) and **Moghania vestita** (baker) syn: Flemingia procumbens

The white root tubers of baker, a leguminous root crop are eaten raw which act as a putative anthelmithic against intestinal parasites. Genistein an active ingredient successfully tested against various trematodes and nematodes has shown good results [64]. Genistein, the major active component which occurs in the edible root-tuber peel of *Flemingia vestita* showed a vermifugal/vermicidal effect on cestodes in vitro by causing a flaccid paralysis and alterations in the tegumental architecture and activity of several enzymes associated with the tegumental interface of the parasites [64]. Tandon et al. [65] reported that genistein obtained from crude extract of baker may influence the glycogen metabolism of *Raillietina echinobothridia*.

**Cucurbita pepo** (Pumpkin)

The pumpkin seeds contain several major groups of active constituents viz. essential fatty acids, amino acids, mucilaginous carbohydrates, phytosterols, minerals and vitamins [49, 66]. Cucurbitacin is a constituent in pumpkin seeds that has shown anti-parasitic activity. Pumpkin seeds have purported effects against tapeworms [66]. The seeds have been shown to effectively treat acute Schistosomiasis in our neighbouring country China.

**Juglans nigra** (black walnut) & **Tanacetum vulgare** (Tansy)

Black walnut and tansy used particularly in treating nematodes infestation especially enterobiosis caused by *Enterobius vermicularis*, a cosmopolitan parasite (Mali and Mehta, 2008) [57]. Presence of tannins, flavonoids and polyphenolic compounds in black walnuts produce anthelmintic activities [67]. The oil of tansy contains tannins and thujone which act against *Nematodirus* spp.

**Mimusops elengi**

An evergreen tree found in the south Asia and Northern Australia commonly known as Spanish cherry, bulletwood. This plant is used as non-astringent, cardiotonic, stomachic and anthelmintic. Tannins of this plant cause uncoupling of oxidative phosphorylation. Stem bark has anthelmintic property [68]. It also contains taraxerol and taraxerene. Bark, flowers and fruits are used in ayurvedic medicine as anthelmintic. Alcoholic extracts showed some efficacy against round worm (*Ascardia goll*) in laboratory studies.

**Punica granatum** (Anar)

Root and stem of the plant is used as an astringent and anthelmintic. Ethanolic extracts of the plant are used in treatment of paraparamphistomosis in infected sheep. It inhibits transformation of egg to larvae of *Haemonchus contortus*. It also shows efficacy against nematodosis. The stem bark is reported to contain an alkaloid, Pelletierine [69, 70].

**Thymus vulgaris**

*Thymus vulgaris* is a species of flowering plant belong to mint family. Thymol an important component has anthelmintic property against *H. contortus*. It shows great efficacy in treating hookworm infection at the dose rate of 2-3 ml extract three times a day [71].

**Kashmir perspective:**

There is an increasing importance in old-fashioned uses of plants for health care among different communities especially in developing countries like India, Pakistan, Nepal, Bangladesh, Sri Lanka etc. Kashmir valley is located in 32°17’ to 36°58’N latitude and 73°26’ to 80°30’ E longitude and an altitude of 1730 m above sea level with Agro-climatic zone of north-west temperate Himalayan region of India having diverse variety of medicinal flora. The traditional uses of different plants against parasitic infections or infestations in both human and livestock is a common practice in the valley. Most of the rural people used the medicinal herbs or plants as veterinary drugs due the expensiveness of the chemical anthelmintics. Based on well-structured questionnaire and detail discussions Tariq and Tantry [72] claimed that 44 plant species belong to 37 genera and 26 families are used as the traditional anthelmints in different preparation and forms which had the anthelmintic properties. Out of these plants, *Artemisia absinthium* (tethwen) and *Achillea millifolium* (pahel-ghassa) were scientifically validated for their claimed anthelmintic action against naturally infected gastrointestinal nematodosis in sheep of the valley [73, 74].

The complete cessation of motility of *Bunostomum trigonocephalum*, the hook worms of sheep after administration of crude aqueous extracts of *Allium sativum* (Rohon) was observed by Khobragade et al. [75] in the valley. Tetrahydro-harmin, the active constituent found in seeds of *Preganum harmala* (Ishband) was found as effective against mixed gastrointestinal nematode infection in goats [23]. The leaves and flowering tops of *Artimisia absinthium* (tethwen) contain volatile oil rich in thujone which has been reported to treat infections with nematodes [76], Lourenco et al, [77] reported that crude aqueous and ethanolic extracts of *Achillea millefolium* (Pahel-gassa) showed significant anthemimtic activity against naturally infected sheep with mixed GI nematodes species. Eugenol, an extract of *Rong* has also been confirmed to possess anthelmintic activity [51]. Crude powder and crude aqueous extract of *Zingiber officinalis* (Adrak) has exhibited anthelmintic activity against gastrointestinal nematodes of sheep [78]. The oral administration of the crude aqueous extracts of *Artemisia absinthium* in sheep was associated with significant reduction in faecal egg output by the GI nematodes [73]. *Chenopodium ambrosioides* (Ganhar) has been reported to cause paralysis and death of worms [72], *Punica granatum* (Daein) and Cucurbita spp. (Aal) have been exclusively employed to treat parasitism due to *Taenia saginata* [72], Wagay [79] in his study from north Kashmir has identified some plants with anthelmintic activity viz. *Angelica glauca* (Chora/Inrin), whole plant extract of *Cardamine macrophylla* (Pahal laish), oil of *Chenopodium urbicum* (Zewa Dawda Kual), seeds of *Portulaca oleracea* (Nunner), leaf extract or paste of *Prunus persica* (Chinan), whole herb extract of *Verbena officinalis* (Hutmool) etc. In Kashmir
valley, Tariq and Tantry [72] conducted a study to observe the anthelmintic properties of some plants and found that root and stem bark of *Acacia arabica* (Kekkar), leaf of *Cannabis sativa* (Bangh), seeds of *Datura stramonium* (Dattur), roots of *Daucus carota* (Gazzer), aerial parts of *Euphorbia royleana* (Guir sochel), fruit and bark of *Ficus carica* (Anjeer), bark and leaf of *Juglans regia* (Doon), whole plant of *Nelumbo nucifera* (Pamposh), whole plant of *Neptea calaria* (Soi), seeds of *Ocimum basilicum* (Babri byol), root and stem bark of *Prunella vulgaris* (Kalaveuth), roots of *Raphanus sativus* (Muj) etc. showed the anthelmintic properties.

**Conclusion**

The advantages of studying local knowledge of herbal medicine are numerous and most of them have already been outlined here. To sum up, it is important, in both a socioeconomic and a scientific perspective, to draw upon the rich store of ethnomedical knowledge. The herbal anthelmintics offer cheaper, more sustainable, available, reliable and familiar alternatives to imported synthetic drugs. Furthermore, when put into production locally, they can reinforce the income and status of local inhabitants. The recognition and development of herbal dewormers offers treatment methods that are more environmentally benign, since they tend to be less toxic, produce fewer unanticipated side-effects, are more biodegradable, accumulate no drug residues in meat or faeces and apparently do not trigger anthelmintic chemo-resistance. However, there are problems connected with the use of botanical dewormer, the largest being the lack of scientific evaluation. Ethnobotanical approach, the most effective knowledge assumes that indigenous uses of plants indicate the presence of biologically active constituents in the plants. This is a slow but very urgent process, as young people desert the traditional knowledge and lifestyles of their elders, leading to the possible disappearance of this knowledge.

**References**


