

Asian Veterinary Medicines: From the Past to the Future

Shuchi GOYAL¹, Divya THIRUMAL¹, Evren ALGIN YAPAR^{2*}, Eda SÖNMEZ GÜRER³, Ashok KUMAR⁴, M Arockia BABU⁵, Rakesh K SINDHU^{6*}

¹ Department of Pharmacognosy, Chitkara College of Pharmacy, Chitkara University, Punjab, 140401, India

² Department of Pharmaceutical Technology, Faculty of Pharmacy, Sivas Cumhuriyet University, 58140, Sivas, Türkiye

³ Department of Pharmacognosy, Faculty of Pharmacy, Sivas Cumhuriyet University, 58140, Sivas, Türkiye

⁴ Department of Cardiology, Sadbhavna Hospital, Fatehabad, Haryana, 125050, India

⁵ Institute of Pharmaceutical Research, GLA University, Mathura 281406, Uttar Pradesh, India

⁶ Department of Pharmacognosy and Natural Products, School of Pharmacy, Sharda University, Greater Noida, Uttar Pradesh, 201306, India

* Correspondence: evren.yapar@yahoo.com (E.A.Y.), rakesh1sindhu16@gmail.com (R.K.S.)

Received: 31 January 2023 / Revised: 25 March 2023 / Accepted: 26 March 2023

ABSTRACT: Asia has plentiful and varied vegetation which is made into active ingredients like probiotics, antimicrobials, and parasiticides are employed to cure or prevent transmission of pathogens as well as to enhance the health of viable cells. World Health Organization researchers have identified over 21,000 plant species that have been used medicinally across the globe. Herbal medicines are risk-free, low-cost, and easy to find. Plants provide 25% of the medications stipulated globally. Almost 75% of pharmaceutical plants grow in nature in various parts of Asia. These plants are known to provide therapeutic effects to various infirmities like poisoning, eczema, osteoporosis, stomachaches, and so on. Veterinary herbal medicines are herbal pharmaceuticals with their remedies, preventive, and indicative uses in zoological cures. Exploring the area of ethnoveterinary medicine is rather unkempt as compared to query on queriesntional cures to diagnose, although growing interest using driven in that prospect. The herbal possessions of Asia also offer the individuals who tend to stock a lush reservoir in contemplating the cure of the animals. Additionally, authenticating and registering the conventional aids and supplies are necessary to ensure it on better to the forthcoming vision. With their recognized molecular processes and potential for further analysis as a prospect for future pharmaceutical drug design and development, traditional medicine/herbal plants are used to treat various diseases in Asian nations.

KEYWORDS: Veterinary herbal medicines; Asian medicinal plants; Ethno-veterinary; Animal-related disorders; Traditional medications.

1. INTRODUCTION

Vata-pitta-kapha are present in every cell, tissue, and organ. They govern psycho-biological changes in the body and physio-pathological changes too. In every person, they differ in permutations and combinations. The balance in the doshas can be affected by (hereditary, congenital, internal, or external trauma, seasonal, natural tendencies or habits, and supernatural factors) [1].

The Chinese emperor Shen Nong compiled and assembled *Materia Medica* around 3700 BC. He was endorsed with savoring various herbal plants and choosing persons that were proper as designating the medical possessions. Due to his strengths, frequent plants turn out to be characteristically expended for therapeutic purposes. His publications of therapeutic herbaceous plants inclined *Materia Medica*. The Shen Nong Jing (Classic of Shen Nong) is a manuscript that was first mentioned by writers who were active in the years after the end of the Han Dynasty (220 AD), who raises the possibility that it was composed during the end of the Han Dynasty. People frequently claim that Chinese medicine has a 5000-year history. However, there is not

How to cite this article: Goyal S, Thirumal D, Algin Yapar E, Sönmez Güre E, Kumar A, Babu MA, Sindhu RK. Asian Veterinary Medicines: From the Past to the Future. *J ResPharm.* 2023; 27(4): 1313-1328.

much evidence available on the usage of herbal remedies prior to the compilation of the Shen Nong herbal, which took place roughly 1800 years ago. "Medicines should coordinate [with one other] in terms of yin and yang, like mother and child, or brothers. To treat cold, one should utilize heat medicinals, to treat the heat, one should use cold medicinals" says Ben Cao. Shen Nong could taste the herbs distinctly and incorporated their qualities into the Tao, the philosophy that served as the foundation for how people saw their environment. Herbs worked with individuals as instruments to move and rebalance their bodies and restore health. The ancient Shen Nong Ben Cao Jing has been released in English translation [2]. Before the Zhou dynasty (1122–770 BC), the early Chinese medical practitioners provided care for both humans and animals [3]. In China, the oldest literature from Shang Dynasty referred to the ailments and cures for horses (1766-1027 BC). Bai Le's Canon of Veterinary Medicine, which Sun Yang penned around 650 AD, was one of the first works in Chinese veterinary medicine. In Mesopotamia, the Sumerians began using cuneiform writing around 3500 BC. The earliest clay tablet from Sumerians still in existence dates to around 2100 BC and lists 250 plant-derived medications in addition to 120 mineral pharmaceuticals and 15 medical prescriptions. "Treatise of Medical Diagnoses and Prognoses," the oldest known medical treatise, dates back to circa 1600 BC. Another well-known text that emerged from Babylonian civilization around 1780 BC, the Code of Hammurabi, addressed how animals were treated, how much it cost, and the consequences of mistreating and making mistakes [4]. In the 1700 BCE, Edwin Smith Papyrus was discovered in Egypt and is now preserved in the New York Academy of Medicine. The circulatory system is described in these scrolls in remarkably realistic detail, with the mention of the heart's fundamental function and the presence of blood arteries all over the body. The Ebers Papyrus, which is currently housed in Leipzig's University Library, dates back to around 1500 BC and contains more than 800 prescriptions. The Library of Alexandria had many more of the old scrolls, but it was burned to the ground in 47 BCE. The Kahun Veterinary Papyrus (dated from circa 1900 BC) [5], on which cattle are prominently shown, is one ancient Egyptian parchment that contains early evidence of veterinary herbal therapy. Compared to Egyptian cultures, ancient Greek and Roman societies started veterinary medical advancements in comparable, but slightly different, ways. One of the first records about Roman practitioners and their research on horses is the "Hippiatrica" [6]. Around 500 BC, the Greek word "Hippiatros" was used to describe horse physicians [4]. Because society's members relied on horses for military and commercial purposes, the horse played a crucial role in Greek and Roman culture. Aristotle, frequently referred to as "the Father of Veterinary Medicine," attained great prominence in Greek culture between 383 and 322 BC. Aristotle wrote on a variety of specialist subjects in his writings, including disease, comparative anatomy, and physiology. Hippocrates had a significant impact on both veterinary and herbal medicine (460-377 BC). He is credited for developing the humoral theory and for writing *Corpus Hippocraticum*, which contained descriptions of more than 200 plants [7].

Veterinary treatment through the earliest eras has not once remained the popular topic everyone is disappearing to investigate, proceedings are often partial, as well as generally patched collected consuming numerous obtains, on the other hand, renditions from ancient languages. TCM eras ago, around 10,000 eons back. Emperor Fusi taught the present primitive Chinese association to domesticate species, according to a folktale. As a result of the requirement to care for domesticated animals, Fusi established veterinary herbal medicines-VHM in China. Shepherds in Middle Eastern nations cared for their animals with a "crude awareness" of basic medical procedures and abilities. Ancient veterinarians employed primitive herbal remedies throughout the Stone Age. Around 3000 BCE, a man by the name of Urlugaledin who lived in Mesopotamia was well-known for his skill in animal healing.

One of the earliest and most comprehensive written legal rules is the Code of Hammurabi. Hammurabi not only outlined veterinarian costs in the Code but also penalties for misconduct. Hippocrates was widely recognized as the founder of medicine and was born in Grecian sunlight in 460 BCE. His work on humoral pathology and the idea that "disease was the product of environmental factors, diet and living habits, rather than a punishment inflicted by the gods" had a long-lasting influence on veterinary and human medicine [8].

Ethno-medical system: A Prominent and Extensive Influence on Veterinary

Herbal medications are one of the ancient practices of remedies renowned and utilized by all ethnic groups and all nations. According to the World Health Organization-WHO, botanic drugs are utilized by 70 percent globally [9], it is hardly surprising that animals, as they have been involved with human existence for a such long time, so man have employed similar plant remedies given to animals under their concern. Ayurveda, India's ethnomedical system, has piqued the curiosity of many individuals and practitioners in recent years. Ayurveda has had an insightful impact on the medication which is practiced in each tradition internationally. Ayurvedic herbs have the earliest ancient usage; their activity after the process and adversarial effects have been assessed via trial and error which allows the specialist a huge use the herbal plants carefully and then,

effectually in veterinary prescriptions. Ashoka constructed one of the first veterinary clinics, on the other hand, simultaneously constructed the functioning rules for veterinary hospitals interpreting the usage of botanic medicines [10]. Ayurvedic medicine traditionally exerted a profound influence beyond Asia, resulting in the establishment of Traditional Chinese Medicine-TCM.

Present-day veterinarians intend that organizations of ethnomedicine are studied the ideologies including customs of TCM by culture to prepare veterinary acupuncture. TCM-skilled animal doctors or practitioners speedily recognize Ayurvedic health ideologies and beliefs. It recalls the physique, observation, and soul and creates an easy link between them. Prana, like Qi, is not gaseous. However, oxygen gas is thought that present in Prana's components. The various locations, hues, and surface coatings have exact analytic explanations. Most Ayurvedic practitioners employ pulse diagnosis. Fingers are aligned in accordance with TCM conformity for the Ayurvedic pulse. Ayurvedic herbal formulations applied with every kind of physical form category are currently made by advanced pharmaceutical businesses in India. The roots of these broad-spectrum compositions are traced in Ayurvedic ancient literature. Indian pharmacological research has focused on achieving an indulgent deed of Ayurvedic botanicals. "Shen Nong" gathered one of the earliest recorded "Materia Medica". The herbal Materia Medica book included herbal products and formulations for both humans and animals. Herbal plants were systems that interacted with people in order to change and stabilize their health [11]. Medications made for animals were segregated as a discipline of traditional Chinese medicine, ancient medical practitioners cured humans and faunas, according to the species [3,12] also in China, the first mention of horse ailments and cures emerged in Shang Dynasty texts (1766- 1027 BC). "Bai-le's Canon of Veterinary Medicine" published by Sun Yang around 650 AD, was one of the first works in Chinese veterinary medicine [13].

"Based on the above potential pharmacological activities of a single herb, the herbal powder seemed not only to improve the uterine condition but also improved the overall physical health of cows with the retained placenta. Additionally, herbal powder treatment could improve the uterine environment due to an earlier placental detachment and a lower puerperal metritis risk. And thus the herbal powder is beneficial for improving the subsequent fertility of cows with retained placenta. Ancient Greek and Roman societies began developments in veterinary medicine in similar, yet slightly different directions compared with the Egyptians. The Hippatrika is one of the first documents we see that relates to Roman practitioners and their study of horses" [6,14]. Kampo talks about the TCM and literally means "the Han Method", referencing to the Chinese herbal method. China and Japan have had traditional links since prehistoric days. Kampo comprehends acupuncture and other TCM components but focuses more on herbal prescriptions. It contrasts with the practice of Chinese herbal medicine in China today notably in its emphasis on the distinctive compendium of herbal formulas. Kampo medicine is extensively followed in Japan [15]. "Rational approaches to medicine were developed to the highest level in the Middle East, where Avicenna lived from 980 to 1037 AD. Abdullah Ibn Ahmad Ibn al-Baytar, as he was known in his land, was a botanist (he described more than 1400 medical herbs, comparing them with the descriptions of ancient authors) and pharmacist. He further codified Galen's theory of using opposites to correct disease processes. His Canon of Medicine is one of the most influential medical books of all time. The Canon described the primary constituents of the human body as the elements – earth, air, fire, and water – that possess two qualities each. The traditional knowledge of the use of medicinal plants which was transmitted from generation to generation is recently in imminent danger of disappearing" [16].

"The culture of Anatolia is a special fusion of Eastern and Western heritage. The historical arteries contain enormous evidence of all the advancements which have been exposed through folklore investigations. Written compositions are named "Baitarname" which are described by local people. These sources state that there are three categories of folk veterinary practice, including magico-religious, empirical, and rational forms [17,18]. Herders, farmers, and rural community members are the only people who understand the ancient medical system. "Some important plants like *Dipsacus inermis*, *Rumex nepalensis*, *Angelica cyclocarpa*, *Saussurea lappa*, *Aesculus indica*, etc. are having great significance in the ethnoveterinary practices." Farmers and nomadic pastoralists are profoundly reliant on therapeutic plants because of the high-cost perspective for operating allopathic veterinary medicines for the fauna. For ages, communes around the world have been using what is now called "traditional systems" to cure and manage parasites and predators that frequently attack cattle, pigs, small ruminants such as goats and sheep, and Indigenous hens [19]. Treatment and control strategies based on medicinal plants are the most widely employed to keep parasites and predators at bay. There is a growing desire for indigenous knowledge systems-IKS to be absorbed and protected since they provide unparalleled advantages to both the growth of Indigenous chickens and consumers [20]. As a result, the usage of ethnoveterinary medicine-EVM, particularly indigenous medicinal herbs, and its associated expertise is

dwindling. The introduction of contemporary technology and restricted knowledge transmission from one generation to the next are frequently identified as primary causes of this escalating problem [21]. Thus, it is critical to preserve and outline local knowledge. It is not unexpected that medicinal plants are being grown to cure and prevent parasites and illnesses that damage indigenous fowl. Sorghum leaves were mashed and given straight to hens. The juice taken from the aloe leaf was combined with water and given to the hens to drink. The ash produced by burning the stems, leaves and bark of *Miora aaxale* was applied directly to the chicken's afflicted region. Indulgent predators, parasites, and therapeutic herbs used to control parasites are critical to making educated decisions while establishing long-term invasions [22] TCM diagnosis and therapy emphasize the intentional aspects of well-being and dysfunction.

Relationship with Human and Zoological Doses

The exigency for herbal medications in veterinary preparation is propagating promptly. Widespread writings and Materia Medica are obtainable on the usage, lethal causes, and discordance in individuals, but the harmlessness of herbal formulations in dogs and cats. For millennia stocks have been associated with herbal plants. Livestock is designed to assimilate and process herbs and herbal plants. The aggressive and addictive properties of the individual herbal formulations are analogous to the dog. Because cats are more delicate, greater attention is required when employing herbal formulations. The compatible conversion of a pharmaceutical dose from one fauna breed to some other, as well as, the rendition of zoological dosage into man dosage is mutually critical after the goal of medicine well-being and efficacy. Furthermore, coercive administration, incorporation, and compositions of medications with feed are commonly used to feed the medicines to fauna. It is recommended that the conversion of zoological doses to man doses be appropriately implemented following regularization to body surface area (BSA) [23]. Ethno-veterinary plant usage is frequently maintained for human use, however, there are additional therapeutic roles in human healthiness rather ethnoveterinary medications (21 percent) In addition, herbs are employed in a variety of ways in zoological and man care (36 percent) or incomplete different methods (38 percent). Specifically, variety of plants, as well as, a wider range of applications are being developed to cure beings rather than animals among Algerian steppe nomads. Concurrent classes occasionally have equivalent use. These substantiate that thoughtful awareness of the medicinal plants and its possessions functions both human beings and other fauna. Regardless of a huge variety of ethno-veterinary ailments, a greater number of plant species are used to treat human health and these are used for better determinations than in ethno-veterinary medicine. Similarly, there is unanimity among local familiarity proprietors in employing ethnoveterinary medicines rather than curing being healthy. When coupled with the circumstances of provided flora, these therapeutic plant enumerates of zoological and human care are expressively related. The logistic regression validates the relationship amidst veterinary medicated herbs and when utilized in man treatment [24]. The efficacy of herbs in a human being is shown in figure1.

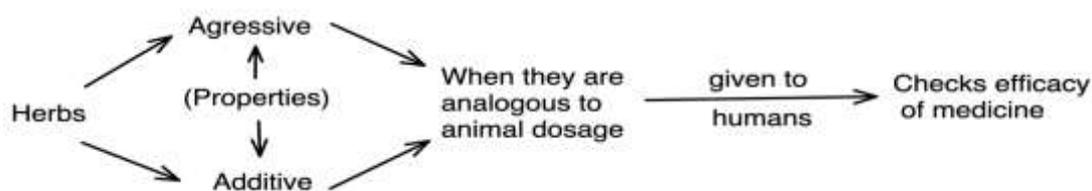


Figure 1. Flowchart shows the properties of herbs given to humans to check the efficacy of Active Pharmaceutical Ingredients-APIs.

2. Ethnoveterinary herbal medicines and their aspects

Flora have been used generally for the remedy and inhibition of various disorders in human and domestic animals. The remedial significance of these herbal formulas varies upon their reactivemodalities that are advanced and sequestered from crude plants, thus emitting a discerning accomplishment on the fauna. Currently, traditional medications are invigorated in veterinary medicine due to their assuring healing efficiency negligible adverse effects of chemotherapeutical agents, and subsiding of drug deposits in animal products that ingested by the individual [25]. An exact organizational attitude in ethnopharmacology consistently necessitates the usage of a list that preferably aids two primary functions: data storage and analytical facilitation, such as quantification and judgements.

“There is an increasing substantiation to explain that synergistic and/or side-effects counteracting combinations of local herbs. Herbal medicine as an alternative remedy has already developed and is likely to play the more significant role. The scientific and local names of mostly used herbs are essentially requested as

they may apply to more than one scientific species, which may or may not be closely related. For example, there are a number of plant species of 'Chamomile' including *Anthemis nobilis* L., *Matricaria chamomilla* L., *Matricaria discoidea* DC, *Cotula matricarioides* (Less.) Bong and *Tanacetum annuum* Pursh. On the contrary, a scientific species may be famous by a number of local plants and classify in folk medicine as they do not correspond to the same botanical category." [26, 27].

Ethno-veterinary experimentation is termed as the organized inspections as well as, utilization of veterinary traditional awareness, thesis, and methods. Ethno-veterinary subjects have been managed chiefly in African, Asian, and American countries [28]. Self-made remedies are frequently used in animal health care, especially where the approach to allopathic drug supplies is indicated or prohibitively overpriced to regional agriculturists [29]. Herbal ethnoveterinary medications are broadly in Himalayas and nearby natives because steer raising became crucial component of the occupation [30]. Conventional herb treatments cater effectual therapies alongside their mutual convenience as compared when to western medications [31]. This traditional awareness is precisely connected compared to adversity in rural populations for generations [32]. The ethnoveterinary awareness for accepting dangerous attrition so that individuals can alter their inclinations is expected to brisk combinations of social and economic alterations around native harmonized respective to the ecological variations and technical progressions [30,33]. It is an intriguing issue to analyze the benefits of this herbal ethnoveterinary that is proportional to rising expenditure going from animal breeding along with care. Moreover, ethnoveterinary medications can be vigorous as well as adaptable so as to cure various categories of stock diseases [34]. Indigenous expertise has consistently been propagated along production deprived of accurate citations as well as conservation [35]. Ethno-veterinary practices have grown sensational prominence over the last ten years due to the new formulations and products of some useful ethnoveterinary products [30,36].

They postulate a cheap assay and minimal expediency contraindicated with modern drugs [31,37]. Currently, those are added reciprocated due to varied socioeconomic factors [38]. Due to its reliance on agriculture and cattle, Pakistan ranks fifth in the world in terms of milk production [39]. Stock farmers in Pakistan have access to almost 4 thousand years of expertise also, experience [40]. The customary comprehension is authorized verbally along one bringing forth to another forth although can be inexistent because of the existing brisk combinations of the social and economic, provisional, along with mechanical developments [41]. Every ethnoveterinary formulation in the analysis section is concocted using herb leaves. The largest utilization of leaf of hefty numerical of ethnomedicinal and ethnoveterinary knowledge are authenticated by various continents [42,43]. Ethno-veterinary medicine, this terminology used for ancient and ethnic zoological protection, implements inexpensive preference to western medications [44]. Human and animal existence have also been circulating all over herbal due to usefulness like feeds, textiles, and shelter, as well as to manage and alleviate illnesses, viz., traditional and consistent repetition [45]. Diseases and herbal medicines differ globally, so, the type, recurrence, along with ministration change due to affinity to natural features, duration, as well as, realization. Ethnically, herbals are utilized to aid disorder, on the other hand, many ailments in a single timing were cut in half since R&D prioritized the use of synthetic compounds. They have been determined to retain herbal products that are element steady, operative, less adverse effects [46]. The ethnoveterinary medicinal plant species are diminishing due to disproportionate and unwise operations, over-foraging, environmental change, population growth, and improper collecting methods such as digging out the entire herbs, advertising strains, and denuding of trees. The herbals are harvested in the highlands, deliver to the provincial marketplaces through locals, and after that sent to metropolitan places. Local workers, partners, as well as plant collecting agents are to be informed of the importance of protecting the site's plant resources, and indigenous people should be included in conservation efforts.

"The ethnoveterinary knowledge of the Kyrgyz people was used successfully until the beginning of the 20th Century. During the Soviet period, many traditional skills, including the treatment, maintenance, and grazing of livestock were centralized, and the prohibition of the practice of ethnoveterinary skills caused a weakened ethnoveterinary knowledge. After the establishment of Kyrgyzstan, farmers again began to openly apply ethnoveterinary methods for the treatment of various livestock ailments. Moreover, scientific studies in this field have begun to be carried out, and a course in ethnoveterinary medicine was opened in some universities in the Kyrgyz Republic."

Role of Phytochemical Characterization of Veterinary Herbal Medicines

Standardization of herbs that are made into medications is required to make sure that more than one of the veterinary herbs medication significant plant constituents and more additives are prevalent in a specified number, as well as to establish their reliability, uniformity, and replicability. Quality standards are also

required for batch-to-batch homogeneity, dose accuracy, durability, and the identification of adulteration. Characterization of physiological phytochemicals in botanic is necessary for quality control-QC, as well as, establishing dosage of herbal-based medications. Because of understanding for a proper dose due to the herbal pharmaceuticals is required, as many plants are effective as veterinary medicines when used in little proportions, but dangerous if used in large amounts [47]. Herbal medications are difficult to standardize since they are made up of complicated mixes of diverse substances. As a result, the plants that have a medical effect are frequently unknown. The standardization and validation of active ingredients require an understanding of the physicochemical qualities of medicinal herbs, as well as additional pre-formulation. Calibration is done using a variety of chemical, spectroscopic, and microbiological approaches [48]. Good manufacturing practices-GMP is a framework that assures in goods are systematically created, that they are managed according to quality requirements, and reduce production hazards that cannot be avoided through finished goods verification. GMP encompasses all areas of manufacturing including QC, from raw resources, facilities, and tools to employee training, safety protocols, and hygienic practices. It also guarantees that suitable standard operation procedures-SOPs to be trailed and accordingly, the task atmosphere is managed, and quality assurance-QA, packing, and labeling for completed complied to specifications. To ensure the quality of veterinary medications, pharmacopeias may consider establishing monographs for herbal compounds which are used exclusively in veterinary science [49,50].

3. Plants Used in Various Disorders

3.1. Herbs Used for Cancer

Herbs that are used for cancer as depicted in the Table 1.

3.2. Herbs for Respiratory Disorders

The liver is a vital body part, which plays an imperative function in modifying numerous biological activities in the form. It is immersed in various essential purposes, like digestion, secretion, nutrient storage, and energy. It has the inordinate capability for detoxication and accretion of endogenous constituents. The liver is probably not only to accomplish biologically operations but also to protect contradicts the risks connected with detrimental doses and compounds. It is generally bared to xenobiotics, hepatotoxins and chemotherapeutic mediators which weaken its purposes [58]. One of the most dangerous illnesses is liver disease. Herbal formulations from plant ingredients have been created as probable liver-protective substances and mediators with a variety of compound structures [59]. Herbs for Respiratory Disorders as shown in Table 2.

Table 1. Herbs for cancer treatment

Herbs	Action	Reference
<i>Astragalus membranaceus</i> L.	Causes cell variation, and apoptosis and shows anti-cancer properties by stimulating toxic mechanisms in cells.	[51]
<i>Withania somnifera</i> L.	Anti-cancer and radiation stimulation properties (efficacious when grouped with radiations).	[52]
<i>Panax ginseng</i> C.A. Meyer	Stimulates differentiation of cells, which lessen the properties of cancer causing agents, moderates inflammation, causes cell death, reduces replication.	[53]
<i>Echinacea purpurea</i> (L.) Moench	Consuming of Echinacea maintains enhancement of Natural killer cells, which are fundamentals in immunosurveillance counter to extemporaneously evolving cancer cells.	[54]
<i>Rumex acetosella</i> L.	This polysaccharide exhibited anticancer ability in mice which are implanted with tumor (180 solid cancer cells).	[55]
<i>Salvia miltiorrhiza</i> L.	It blocks DNA formation and tumor proliferation; gene expression; blockage due to the telomerase ability of tumor cells.	[56]
<i>Piper methysticum</i> L.	Flavokavains shows potent antiproliferative and cell death activity in bladder tumor cells.	[57]

Table 2. Herbs for Respiratory Disorders

Herbs	Action	Reference
<i>Cissampelos pareira</i> L.	The treatment with ethanolic root extract of <i>Cissampelos pareira</i> L. shown a strong protective effect by reducing blood biochemical indicators. The administration of root extract with CCl ₄ produced a protective effect on the liver.	[60]
<i>Citrus reticulata</i> L.	All metabolic markers were significantly reduced in mice treated with essential oil of <i>Citrus reticulata</i> L. <i>Citrus reticulata</i> L. has strong hepatoprotective effect against isoniazid-induced liver injury.	[61]
<i>Tecomella undulate</i> Seem	It was discovered that the hepatoprotection provided by <i>Tecomella undulate</i> leaves against liver damage might due to quinones, along with antioxidants.	[62]
<i>Benincasa hispida</i> L.	Aqueous extract of <i>Benincasa hispida</i> L. pulps has a high concentration of antioxidants, which may protect against diclofenac-induced hepatotoxicity.	[63]
<i>Phyllanthus polyphyllus</i> L.	<i>Phyllanthus polyphyllus</i> L., a herbal plant, has been proven to have potent hepatoprotective and antioxidant properties.	[64]
<i>Calotropis gigantea</i> L. Dryand. ex W.T. Aiton	The extract's pharmacological activity may be attributed to its antioxidant potential to suppress lipid peroxidation as well as prevent vitamin C depletion.	[65]
<i>Orthosiphon stamineus</i> Benth.	<i>Orthosiphon stamineus</i> is an important Ayurvedic herb that has been shown to protect the liver against thioacetamide-induced hepatotoxicity in rats.	[66]
<i>Aerva lanata</i> L. Juss. Ex Schult.	When administered with a hydro-alcoholic extract of <i>Aerva lanata</i> , blood biochemical parameters such as bilirubin and were considerably changed, proving it to be a hepatoprotective herbal medicine.	[67]
<i>Allium cepa</i> L.	<i>Allium cepa</i> L. shows hepatoprotection against Cd induced liver damage and cures liver disorders.	[68]

3.3. Herbs for Dermatologic Disorders

Herbal formulations and sequestered plant mixtures recreation a rising part of the therapy of skin diseases and lesions. The healing potential of medicinal herbs historically used in dermatology has been investigated in recent years, and some of them have been produced and authorized as drugs or medical devices for the treatment of skin problems. Furthermore, a growing number of herbal products, sometimes referred to as "cosmeceuticals," have been produced in the field of medical cosmetics. Herbs for Dermatologic Disorders as shown in Table 3.

Table 3. Herbs for Dermatologic Disorders

Herbs	Action	Reference
<i>Hypericum perforatum</i> L.	It has antibacterial, anti-inflammatory & keratinocyte differentiation-promoting effects and is used to treat wounds and burns.	[69]
<i>Glycyrrhiza glabra</i> L.	It has an anti-inflammatory effect.	[70]
<i>Potentilla erecta</i> (L.) Raeusch	It treats eczema & has anti-inflammatory effects.	[71]
<i>Vataireopsis araroba</i>	Effective psoriasis therapy, blocks the pro-inflammatory cytokines as well as keratinocyte growth.	[72]
<i>Salvia officinalis</i> L. & <i>Rheum palmatum</i> L. (Combination)	In individuals with herpes simplex labialis, this medication is used (causes inflammations in some parts of the animal dermal part)	[73]
<i>Euphorbia peplus</i> L.	The principal pharmacological considerations stages the actinic keratosis is a differentiation-promoting impact.	[74]
<i>Betula</i> spp.	It influences 3 stages of injury (the inflammation & shift & differentiating process of cells),	[75]

3.4. Herbs for Neurologic and Behavioral Disorders

Herbal remedies have been utilized to treat neurological disorders since ancient times. Chronic inflammatory reactions in the neurological system have a degenerative effect on the central nervous system-CNS, and herbs and their chemical contents operate as powerful neuroprotective agents. A nervine is any plant that affects the nervous system in traditional herbal therapy. As a result, a nervine might comprise a variety of therapeutic activities. These relaxants have a sedative, hypnotic, relaxing and anxiolytic effects, and many have antispasmodic activity. Anxiety, hyperactivity, insomnia, and restlessness are all treated with relaxant nerviness. Activation can advantageous in neurodegeneration or less active neuronal system, and nervine tonics can directly rejuvenate tissues by helping injury which further assists minimize efficient unnecessary strain. Herbs for Neurologic and Behavioral Disorders as depicted in Table 4.

Table 4. Herbs for Neurologic and Behavioral Disorders

Herbs	Pharmacological Action	Reference
<i>Crocus sativus</i> L.	In humans and animal models, it possesses anticonvulsant and anti-Alzheimer action. Treating depression and influencing Ach application & synergy along alkaloid category.	[76]
<i>Coriandrum sativum</i> L.	Treat insomnia, reduce time for active convulsions & shows its action.	[77]
<i>Ocimum sanctum</i> L.	In acute seizure models, it has a neuromodulatory & anticonvulsant impact. It protects and stimulates nerves while also preventing Reactive Oxygen Species-ROS activation, DNA repaired, & cell depolarization.	[78]
<i>Nigella sativa</i> L.	Anti-oxidative action, works as a neuroprotective agent against a variety of nervous system illnesses including Alzheimer's disease, epilepsy, and neurotoxicity.	[79]
<i>Ferula asafetida</i> L.	It has flatulence relieving, anti-helminthic, increases perspiration qualities, also possesses anti-pyretic action. It is also cures cough, relieves pain, antiperiodic, antidiabetic, anti-spasmodic, anti-inflammatory, contraceptive, and antiepileptic.	[80]
<i>Panax ginseng</i> C.A. Meyer	It prevents neurodegeneration, increases cholinergic metabolic rate, and suppresses apoptosis and neuronal loss	[81]
<i>Corus calamus</i> L.	It improves learning performance and behavior and works as a neuroprotective for Parkinson's disease in mice.	[82]
<i>Centella asiatica</i> L. Urb.	It is utilized as a psychotropic therapeutic herb, used to relieve oxidative stress symptoms, used to increase remembrance by model.	[83]
<i>Salvia officinalis</i> L.	Memory-enhancing herb with anti-inflammation activities & anti-oxidant factors, also as a modest prohibition of the action on AChE.	[84]

3.5. Herbs for Cardiovascular Disorders

The current surge of interest in alternative medicine and natural goods has reignited interest in ancient therapies used to treat cardiovascular conditions. Patients with congestive heart failure-CHF, systolic hypertension, angina pectoris, atherosclerosis, cerebral insufficiency, venous insufficiency, and arrhythmia have all been treated using native herbs. Several botanicals contain powerful cardioprotective agents, which exert cardiac effects on the heart. Herbs for cardiovascular disorders are shown in Table 5.

Table 5. Herbs for cardiovascular disorders

Herbs	Pharmacological Action	Reference
<i>Allium sativum</i> L.	Inhibit the pathogenesis of cardiovascular disease (increasing blood total cholesterol, increased LDL (low-density lipoprotein) oxidation, increased platelet aggregation, hypertension).	[85]
<i>Commiphora wightii</i> (Arn.) Bhandari	Prevents obesity, hyperlipidaemic, hypercholesterolaemic, acrid & septic, arthritis, microbial, inflammation & cancer.	[86]
<i>Crataegus oxyacantha</i> L.	It cures heart diseases like pectoris, high BP, heart attacks, Congestive heart failure. Elevated myocardial contraction & decreases other parts vascular resistance, which increases efficacy of glycoside also, contains some anti-hypertensive factors.	[7]
<i>Terminalia arjuna</i> L.	Antidysenteric, relief from chest discomfort, useful in lowering BP and HR in animals. Useful in treatment for CHF, high BP & blood deficiency causing stroke.	[88]
<i>Plectranthus barbatus</i> Andr.	It elevates myocardial muscle & other part vasodilating which reduces heart pre-load & after-load,	[89]
<i>Sanguinaria canadensis</i> L.	Enhance contractility and provide a compact-dependent increasing muscle contractility.	[90]
<i>Zanthoxylum americanum</i> Mill.	Usage to describe "sluggish circulation." (Cardiac activity is intensified, the pulse increases somewhat faster, and the integumentary glands secrete abundantly).	[91]
<i>Salvia miltiorrhiza</i> L.	ACE inhibitory activity in a hypertensive rat, used to treat blood stagnation and chest discomfort, decreases platelet activity and increases fibrinolysis.	(92)

4. Use of Herbal Feed Additives for Improved Animal Production

The exploitation of artificial medicines utilized for animal rearing indicates transmission in medicines resistance-showing bacteria for people & accumulation of lethal remains in meat and milk. Utilization of herb feed additives that attain significance in living animals' production, with the suitable so that it reduces the usage in significant type medicine, damaging enduring properties also, price efficacy. The variety of feed additives including probiotic-microorganism (health boosting the microorganism), prebiotics-ingredients (fermented allows specific changes), and acerbic chemicals as well as, herb active constituents show demonstrated to improve lasting characteristics to improve animal productivity. These plants include a variety of bioactive chemicals that are typically considered biodegradable. Typically, mixtures allow herbs engage to in their surroundings also, operate as a defense arrangement in contrast to biological and developmental stress, as well as predators or infections. Aside from hazardous chemicals, numerous of these secondary plant metabolites have been shown to have positive impacts on food items and animal metabolism. Herbs or phytochemicals can selectively alter bacteria by antimicrobial action or by stimulating eubiotics of the microbiota [93]. A significant proportion of herbal feed additives exhibit their antimicrobial property by polymerizing and congealing antigens in the bacteria cell wall composition. The natural oils change the cytosolic membrane permeability to H⁺ and K⁺ ions. This modification affects critical cellular activities including like transfer of electrons, and proteins when synthesized in a cell across the membrane through the complex, link of process to the electron transport chain, also the catalyst-requiring events, leading to a decrease of biological ion regulation & as a result, bacteria mortality [94]. The lipophilic property of hydrophobic liquid concentrates certain collect in the layer of cells causing rupture of the bacterial cytoplasmic membrane. Other actions may be related to nutrition absorption inhibition, enzymatic inhibition, DNA, RNA, and protein production by bacteria. Inhibition of oxidation is mostly phenol-like chemical requirements, flavonoid-like constituents & terpenoid-like constituents, which protect collection cells from harmful effects in auto-oxidation processes. The viable methods of action of herbal plants in the physical progress preferment contain variations that show gut microorganism genetic material, the elevation of nutrient absorbed by the body,

increase nitrogen processed, showing performance of the high degree of resistance to an illness, physiological & tissue alterations of the GIT. Finally, plants contribute to the nutrition requirements of the fauna species. The herbal feed additives utilize their favorable impacts:

Antimicrobial- Herbal plants wield anti-microbial mechanisms in-vitro-lab conditions in contra to imperative pathogens. They employ their motion-concluded altered processes. These mechanisms ascend primarily which shows the perspective in the active agents as intervene in bacteria's layer of outer cells, disorganizing the cells layer and its structures & causing ionic exudations [95]. Tannins are metabolite ingredients implicated in the action of herbal feed additives functioned through ferrous detriment, H-bonding or non-selective interactions with a critical organic molecule like a catalyst [96]. The capacity of an alkaloid to impede DNA synthesis is the mechanism through which it exhibits antimicrobial actions. Along with other supporting herbs, these plants include important flavonoid components such as baicalein, limonene, cinnamaldehyde, carvacrol and eugenol having antibacterial properties [97].

Anti-inflammatory- Curcuma, black pepper, cloves, cinnamon, mint, and ginger extracts were shown to be anti-inflammatory. Phenols, terpenoids, and flavonoids are the main anti-inflammatory active chemicals. These molecules prevent the breakdown of inflammatory prostaglandins [98].

Antioxidant- Mechanism is shows the result primarily through ions scavenging and modulating the mechanism for antioxidant catalyst [99].

Immunostimulant- Polysaccharide is a primary immune active ingredient present in Chinese herbs, and as such, it is recognized as the cornerstone of increasing immune function. Plants that contain immunostimulatory compounds include echinacea, licorice, and garlic. Plants elevate the active action of lymphocytes, destroy the target cells, and destroy the virus, as well as promote phagocytosis and interferon production (100).

Potential merits of herbal feed additives over other additives include:

1. Natural feed component.
2. The absence of any lasting effects.
3. Non-hazardous and environmentally friendly.
4. Minimal drug resistance problem.

5. Herbal Toxicity in Fauna

Because therapeutic effectiveness occurs at a low dosage, an overdose might cause toxicity, and the intercalating issue between pharmacology and toxicology is crucial. Toxic plants, on the other hand, may contain active chemicals with biologically beneficial properties. To minimize toxicity issues and animal death, it is crucial to be mindful of the poisonous possibilities of veterinary herbs [101]. Plant poisoning is closely tied as utilization in hazardous dosages of healthcare, along the several examples, containing death-causing ones, assumed to have occurred without diagnosis or record [102]. In several nations, toxic plant adulteration of human meals and unintentional exposure to plant poisons have been recorded. Furthermore, notably in arid locations, animals may feed on hazardous wild plants. The American Herbal Products Association-AHPA presented a four-class categorization of therapeutic herbs in 1997: the 1st category including herbs with a wide safety, such as Calendula, Hawthorn, Euphrasy, Lavender, Taraxacum, Nettle, Chamomille, Echinacea & Peppermint; the 2nd category, which is divided into 4 categories, includes plants with little limitations including SJW, Sage, Artemisia & Liquorice. The 3rd category comprises plants where the presence of definitive proof that requires the oversight of a professional practitioner; lastly, the fourth class includes all plants that have never been categorized [103]. Combination therapy between herbal and synthetic medications, as well as correspondingly utilized herbal remedies albeit they are uncommon. Other particular limitations might include already made ailments (for instance, gastric ulceration, renal, and liver cirrhosis) else, surgical that alters kinetics including the active ingredients, potentially increasing the risk of adverse consequences. Concluding, an element of the product is critical for assurance of herbs made into pharmaceuticals; which is actually, remnants of environmental contaminants (metallic elements, toxins by some fungus, and unstable form of the elements) in the plant-therapeutic formulations may produce undesirable consequences [104]. *Mentha piperita* L. oil is now utilized in domestic animal medicine as a pest control. Pulegone is hepatotoxic to rabbits and can cause cerebellar lesions in mice rate of dosage of 200milligrams per kilogram body mass [105]. Herbal remedies have already been found in many cases in therapy of a certain fauna, still, human trial are necessary so that it can be usage in another species.

6. CONCLUSION AND FUTURE PERSPECTIVES

The methodologies should be explored to establish the theoretical and clinical foundation for selecting the optimum isolated bioactive components from indigenous medicinal literature. Medicinal herbs have been found for the therapy of a particular breed in many cases; however clinical studies should be necessary for usage in other species. In addition, approaches are formulated so that resolve the experimental as well as efficient category of choice the utmost adequate apportioned reactive formulae which include ethnoveterinary medications composition. Largely happening instances, herbs cure is consistent for the analysis of the critical fauna, on the other hand, human trials are done so that they are appropriate to the utilization in another animal.

The ethnic awareness of the utilization of herbal that is handed-down derived propagation to next is afresh in immediate or in temporary. Therefore, ethnic awareness potency is a choice for further conceptions in the therapy of animal-rear disorders so that it doesn't disappear. Recent studies say an essential improvement against the conservation of native herbal-based information from exterminating. Current ethnoveterinary usage in the studies is established to gastrointestinal toxicity, not hydrated does not get digested, worms are removed, and other instances. Phytonutrients and their mechanism of action experimentations are done so that isolation of the active metabolite, as well as detection of elements in lab conditions or in animal testing efficient with the specified herbs for the addressed animal disorders, are critical. Ethno-medicinal plant knowledge has been passed down through the ages, but it is now in jeopardy of extinction. The recent research makes a significant contribution to the preservation of indigenous plant-based knowledge. To preserve this information, young people should be encouraged to become interested in ethnomedicinal practices. EVM therapies, notably for viral infections, are awaiting more investigation outside of the lab or in publications. This describes the numerous types of herbal remedies and how they function, as well as whether veterinary herbal therapy has potential. As a result, traditional knowledge may be an alternative for developing approaches in cattle therapeutic strategies and, as a result, should be recorded before it gets destroyed.

Abbreviations:

AHPA- American Herbal Products Association
CHF- Congestive heart failure
EVM- Ethno veterinary medicines
GMP- Good Manufacturing Practices
LDL- Low-density lipoprotein
QA- Quality assurance
QC- Quality control
TCM- Traditional Chinese Medicine

Author contributions: Concept – R.K.S., E.A.Y.; Design – R.K.S., E.A.Y.; Supervision – R.K.S.; Literature Search – S.G., D.T., E.S.G.; Writing – S.G., D.T., A.K., R.K.S., E.A.Y.; Critical Reviews – R.K.S., E.A.Y M.A.B.

Conflict of interest statement: The authors declared no conflict of interest.

Acknowledgement: All authors are grateful to Sharda University, Sivas Cumhuriyet University and Chitkara University, for providing institutional facilities.

REFERENCES

- [1] Ekor M. The growing use of herbal medicines: issues relating to adverse reactions and challenges in monitoring safety. *Front Pharmacol.* 2014; 4:177. <https://doi.org/10.3389/fphar.2013.00177>.
- [2] Wynn SG, Fougère BJ. The Roots of Veterinary Botanical Medicine. In: *Veterinary Herbal Medicine*, Elsevier Health Sciences, 2007; 33-49. <https://doi.org/10.1016/B978-0-323-02998-8.50008-1>.
- [3] Schoen A. *Veterinary Acupuncture: Ancient art to the modern medicine*, Mosby, Sydney, Australia, 1994.
- [4] Swabe J. *Animals, disease and Human Society: Human-Animal relations and the rise of veterinary medicine*, Routledge, New York 1999.
- [5] Karasszon D. *A concise History of Veterinary medicine*. Budapest Akademiai Kiado, 1998.
- [6] Walker RE. *Ars Veterinaria: The Veterinary Art from Antiquity to the End of the XIXth Century: Historical Essay*. Schering-Plough Animal Health, Kenilworth, NJ, 1991.
- [7] Wynn SG, Fougère BJ. *Veterinary Herbal Medicine: A Systems-Based Approach*. *Veterinary Herbal Medicine*. 2007:291-409. <https://doi.org/10.1016%2FB978-0-323-02998-8.50024-X>.
- [8] Mark Joshua J. *A Brief History of Veterinary Medicine*. *World History Encyclopedia*. Last modified April 30, 2020. Available at: <https://www.worldhistory.org/article/1549/a-brief-history-of-veterinary-medicine/>

- [9] Eisenberg DM, Davis RB, Ettner SL, Appel S, Wilkey S, Van Rompay M, Kessler RC. Trends in alternative medicine use in the United States, 1990-1997: results of a follow-up national survey. *JAMA*. 1998;280(18):1569-1575. <https://doi.org/10.1001/jama.280.18.1569>.
- [10] Anjaria J. Traditional System Vet Med for Small Farmers in India. RAPA No. 80. 1984.
- [11] The Divine Farmer's Materia Medica: A Translation of the Shen Nong Ben Cao (Blue Poppy's Great Masters Series). Shou-Zhong Yang (Editor). Blue Poppy Press, 1998.
- [12] Haltrecht H. Veterinary Acupuncture: Ancient Art to Modern Medicine. *Can Vet J*. 1995;36(10):646.
- [13] Gao X, Wang W, Wei S, Li W. Review of pharmacological effects of *Glycyrrhiza uralensis* and its bioactive compounds. *Zhongguo Zhong Yao Za Zhi*. 2009;34(21):2695-700. Chinese.
- [14] Marsden, Steve, and Wynn, Susan G.. *Manual of Natural Veterinary Medicine: Science and Tradition*. United Kingdom, Mosby, 2003, UK.
- [15] Dharmananda S. *Kampo medicine, the practice of Chinese herbal medicine in Japan*. ITM, 2002.
- [16] Anyinam C. Ecology and ethnomedicine: exploring links between current environmental crisis and indigenous medical practices. *Soc Sci Med*. 1995; 40:321-329. [https://doi.org/10.1016/0277-9536\(94\)e0098-d](https://doi.org/10.1016/0277-9536(94)e0098-d).
- [17] Başağaç Gül RT, Genc SV. Some examples of animal healing in folk veterinary medicine in Anatolia 15th Annual Conference DVG 'Animal Healers' Past and Present, Berlin, Germany, 2009.
- [18] History of Medicine Pharmacology Veterinary Medicine in Anatolia and Turkic cultures. Sari N (editor). International Congress of the History of Medicine 2002, Istanbul, Turkey.
- [19] Rafique Khan SM, Akhter T, Hussain M. Ethno-veterinary practice for the treatment of animal diseases in Neelum Valley, Kashmir Himalaya, Pakistan. *PLoS One*. 2021;16(4):e0250114. <https://doi.org/10.1371/journal.pone.0250114>.
- [20] Sharma M, Kaushik RP. Plant genetic resources and indigenous traditional knowledge conservation toward resilience to climate change. In: *Plant genetic resources and traditional knowledge for food security*. Salgotra R, Gupta BB (editors). Springer, Dordrecht, 2015; pp.199-214.
- [21] Kiyafat U, Khan F, Shah M, Ahmad H, Ashraf M, Rahman I, Iqbal Z, Mulk Khan S, Majid A. Investigation of traditional veterinary phytomedicines used in Deosai Plateau, Pakistan. *Global Veterinaria*. 2015; 15(4):381-388. <http://dx.doi.org/10.5829/idosi.gv.2015.15.04.96104>.
- [22] Ndlovu W, Mwale M, Iwara IO, Kabiti HM, Obadire OS, Francis J. Profiling village chickens predators, parasites and medicinal plants used to control the parasites. *Brazil J Poult Sci*. 2021; 23(2): eRBCA-2019-1023. <https://doi.org/10.1590/1806-9061-2019-1023>.
- [23] Guidance for Industry. Estimating the safe starting dose in clinical trials for therapeutics in adult healthy volunteers. Department of Health and Human Services, Center for Drug Evaluation and Research, Center for Biologics Evaluation and Research. 2002.
- [24] Maire R. *Flore de l'Afrique du Nord (Maroc, Algérie, Tunisie, Tripolitaine, Cyrénaïque et Sahara)*. Le Chevalier, Paris, 1959.
- [25] Gurib-Fakim A. Medicinal plants: traditions of yesterday and drugs of tomorrow. *Mol Aspects Med* 2006; 27:1-93. <https://doi.org/10.1016/j.mam.2005.07.008>.
- [26] Heinrich M, Edwards S, Moerman DE, Leonti M. Ethnopharmacological field studies: a critical assessment of their conceptual basis and methods. *J Ethnopharmacol*. 2009; 124:1-17. <https://doi.org/10.1016/j.jep.2009.03.043>.
- [27] Pardo-de-Santayana M, Morales R. Chamomiles in Spain: The Dynamics of Plant Nomenclature. In: *Ethnobotany in the New Europe: People, Health and Wild Plant Resources*, 2010, Berghahn Press, pp:282-306.
- [28] Pieroni A, Howard P, Volpato G, Santoro RF: Natural remedies and nutraceuticals used in ethnoveterinary practices in inland southern Italy. *Vet Res Commun*. 2004; 28:55-80. <https://doi.org/10.1023/b:verc.000009535.96676.eb>.
- [29] Nyamanga P, Suda C, Aagaard-Hansen J: The socio-cultural context and practical implications of ethnoveterinary medical pluralism in western Kenya. *Agric Hum Values* 2008; 25: 513-527. <https://doi.org/10.1007/s10460-008-9141-1>.
- [30] Lans C, Turner N, Khan T, Brauer G, Boepple W. Ethnoveterinary medicines used for ruminants in British Columbia, Canada. *J Ethnobiol Ethnomed*. 2007;3:11. <https://doi.org/10.1186/1746-4269-3-11>.
- [31] Ganesan S, Chandhirasekaran Mand Selvaraj A. Ethno-veterinary healthcare practices in southern districts of Tamil Nadu. *Indian J Tradit Know*. 2008; 7(2): 347-354.
- [32] Sullivan K, Shealy CN. *Complete Natural Home Remedies*. Element Books Limited, Shaftesbury, UK, 1997, pp. 3-5.
- [33] Lans C. Possible similarities between the folk medicine historically used by First Nations and American Indians in North America and the ethnoveterinary knowledge currently used in British Columbia, Canada. *J Ethnopharmacol*. 2016 Nov 4;192:53-66. doi: 10.1016/j.jep.2016.07.004. Epub 2016 Jul 6. PMID: 27394389.
- [34] Ullah M, Khan MU, Mahmood A, Malik RN, Hussain M, Wazir SM, Daud M, Shinwari ZK. An ethnobotanical survey of indigenous medicinal plants in Wana district south Waziristan agency, Pakistan. *J Ethnopharmacol*. 2013;150(3):918-924.
- [35] Bullitta S, Re GA, Manunta MDI, Piluzza G. Traditional knowledge about plant, animal, and mineral-based remedies to treat cattle, pigs, horses, and other domestic animals in the Mediterranean island of Sardinia. *J Ethnobiol Ethnomed*. 2018;14(1):50. <https://doi.org/10.1186/s13002-018-0250-7>.
- [36] Rehman S, Iqbal Z, Qureshi R, Rahman IU, Sakhi S, Khan I, Hashem A, Al-Arjani AF, Almutairi KF, Abd Allah EF, Ali N, Khan MA, Ijaz F. Ethnoveterinary Practices of Medicinal Plants Among Tribes of Tribal District of North Waziristan, Khyber Pakhtunkhwa, Pakistan. *Front Vet Sci*. 2022 Mar 25;9:815294. doi: 10.3389/fvets.2022.815294. PMID: 35400104; PMCID: PMC8990162.

- [37] Sharma R, Manhas RK, Magotra R. Ethnoveterinary remedies of diseases among milk yielding animals in Kathua, Jammu and Kashmir, India. *J Ethnopharmacol.* 2012 May 7;141(1):265-72. doi: 10.1016/j.jep.2012.02.027. Epub 2012 Feb 22. PMID: 22366093.
- [38] Rahaman CH, Ghosh A, Mandal S. Studies on the ethnoveterinary medicinal plants used by the tribals of Birbhum district, West Bengal. *J Econ Taxon Bot.* 2009; 33: 333-338.
- [39] Hassan HU, Murad W, Tariq A, Ahmad A. Ethnoveterinary study of medicinal plants in Malakand Valley, District Dir (Lower), Khyber Pakhtunkhwa, Pakistan. *Irish Vet J.* 2014; 67: 6. <https://doi.org/10.1186/2046-0481-67-6>.
- [40] Farooq Z, Iqbal Z, Mushtaq S, Muhammad G, Iqbal MZ, Arshad M. Ethnoveterinary practices for the treatment of parasitic diseases in livestock in Cholistan desert (Pakistan). *J Ethnopharmacol.* 2008;118(2):213-219. <https://doi.org/10.1016/j.jep.2008.03.015>.
- [41] Hussain A, Khan MN, Iqbal Z, Sajid MS. An account of the botanical anthelmintics used in traditional veterinary practices in Sahiwal district of Punjab, Pakistan. *J Ethnopharmacol.* 2008;119(1):185-190. <https://doi.org/10.1016/j.jep.2008.06.034>.
- [42] Kala CP. Ethnomedicinal botany of the Apatani in the Eastern Himalayan region of India. *J Ethnobiol Ethnomed.* 2005; 1:11. <https://doi.org/10.1186/1746-4269-1-11>.
- [43] Bhat JA, Kumar M, Bussmann RW. Ecological status and traditional knowledge of medicinal plants in Kedarnath Wildlife Sanctuary of Garhwal Himalaya, India. *J Ethnobiol Ethnomed.* 2013;9:1. <https://doi.org/10.1186/1746-4269-9-1>.
- [44] Guèye EF. Ethno-veterinary medicine against poultry diseases in African villages. *Worlds Poult Sci J.* 2019; 55:187-198. <https://doi.org/10.1079/WPS19990013>.
- [45] McCorkle CM. An introduction to ethno-veterinary research and development. *J Ethnobiol.* 1986; 6:129-149.
- [46] Arshad M, Rao AR. Medicinal plants of Pakistan of Cholistan Desert. *Medicinal plants of Pakistan. Proceeding of the meeting held at the PGRI, NARC, Islamabad, 1998, PP.1-11.*
- [47] Wani MS, Polshettiwar SA, Parakh SR, Chopade VV. Herbal medicines and its standardization. *Pharm Rev.* 2007; 5.
- [48] Chakravarthy BK. Standardization of herbal products. *Indian J Nat Prod.* 1993; 9:23-26.
- [49] Sane RT. Standardization, quality control and GMP for herbal drugs. *Indian Drugs.* 2002; 39:184-190
- [50] *Veterinary Herbal Pharmacopoeia*, Ed: Sun-Chong Wang, Taiwan, Publication Date: July 2020, DOI: 10.52305/GHTR1903, ISBN: 978-1-53617-947-7.
- [51] Kurashige S, Akuzawa Y, Endo F. Effects of astragali radix extract on carcinogenesis, cytokine production and cytotoxicity in mice treated with carcinogen, N-butyl-N'-butanonitrosoamine. *Cancer Invest.* 1999; 17:30-35. <https://doi.org/10.1080/07357909909011714>.
- [52] Sharada AC, Solomon FE, Devi PU, Udupa N, Srinivasan KK. Antitumor and radiosensitizing effects of withaferin A on mouse Ehrlich ascites carcinoma in vivo. *Acta Oncol.* 1996; 35:95-100. <https://doi.org/10.3109/02841869609098486>.
- [53] Helms S. Cancer prevention and therapeutics: *Panax ginseng*. *Alt Med Rev.* 2004; 9:259-274.
- [54] Brousseau M, Miller SC. Enhancement of natural killer cells and increased survival of aging mice fed daily Echinacea root extract from youth. *Biogerontology.* 2005; 6:157-163. <https://doi.org/10.1007/s10522-005-7951-8>.
- [55] Ito H. Effects of the antitumor agents from various natural sources on drug-metabolizing system, phagocytic activity and complement system in sarcoma 180-bearing mice. *Jpn J Pharmacol.* 1986; 40:435-443. <https://doi.org/10.1254/jjp.40.435>.
- [56] Yuan SL, Wang XJ, Wei YQ. Anticancer effect of tanshinone and its mechanisms. *Ai Zheng.* 2003; 22(12): 1363-1366.
- [57] Zi X, Simoneau AR. Flavokawain A, a novel chalcone from kava extract, induces apoptosis in bladder cancer cells by involvement of Bax protein-dependent and mitochondria-dependent apoptotic pathway and suppresses tumor growth in mice. *Cancer Res.* 2005; 65:3479-3486. <https://doi.org/10.1158/0008-5472.can-04-3803>.
- [58] Pal A, Banerjee B, Banerjee T, Masih M, Pal K. Hepatoprotective activity of *Chenopodium album* linn. Plant against paracetamol induced hepatic injury in rats. *Int J Pharm Pharm Sci.* 2011; 3(3): 55-57.
- [59] Thyagarajan SP, Jayaram S, Gopalakrishnan V, Hari R, Jeyakumar P, Sripathi MS. Herbal medicines for liver diseases in India. *J Gastroenterol Hepatol.* 2002;17 Suppl 3:S370-376. <https://doi.org/10.1046/j.1440-1746.17.s3.30.x>.
- [60] Sing Yadav M, Kumar A, Sing A, Sharma US, Sutar N. Phytochemical investigation and Hepatoprotective activity of *Cissaampelos pareira* Carbon tetrachloride induced hepatotoxicity. *Asian J Pharm Health Sci.* 2011; 1(3):106-110.
- [61] Kangralkar VA, Gavimath CC, Vijapur V, Gowri B, Vinay R, Mathapati HP. Protective effect of essential oil of *Citrus reticulata* on isoniazid induced hepatotoxicity in wistar rats. *Int J Pharm Applicat.* 2010; 1(2):59-61.
- [62] Singh D, and Gupta RS. Hepatoprotective activity of methanol extract of *Tecomella undulate* against alcohol and paracetamol induced hepatotoxicity in rats. *Life Sci Med Res.* 2011; 26:1-8
- [63] Das SK, Roy C. The protective role of *Benincasa hispida* on diclofenac sodium induced hepatotoxicity in albino rat model. *Int J Pharm Res Develop.* 2012; 3(11): 171 - 179.
- [64] Raj Kapoor B, Venugopal Y, Anbu J, Harikrishnan N, Gobinath M, Ravichandran V. Protective effect of *Phyllanthus polyphyllus* on acetaminophen induced hepatotoxicity in rats. *Pak J Pharm Sci.* 2008; 21(1): 57-62.

- [65] Kshirsagar A, Purnima A. Evaluation of *Calotropis gigantea* r. br. flower extract on alcohol induced hepatotoxicity. *J Cell Tissue Res.* 2008; 8(3):1551-1556.
- [66] Alshawsh MA, Abdulla MA, Ismail S, Amin ZA. Hepatoprotective effects of *Orthosiphon stamineus* extract on thioacetamide-induced liver cirrhosis in rats. *Evid Based Complement Alternat Med.* 2011;2011:103039. <https://doi.org/10.1155/2011/103039>.
- [67] Manokaran S, Jaswanth A, Sengottuvelu S, Nandhakumar J, Duraisamy R, Karthikeyan D, Mallegaswari R. Hepatoprotective activity of *Aerva lanata* Linn. against paracetamol induced hepatotoxicity in rats. *Res J Pharm Techn.* 2008; 1(4):398-400
- [68] Ige SF, Akhigbe RE, Edeogho O, Ajao FO, Owolabi OQ, Oyekunle OS, Ajayi AF. Hepatoprotective activities of *Allium cepa* in cadmium-treated rats. *Int J Pharm Pharm Sci.* 2011; 3(5):60-63.
- [69] Schempp CM, Windeck T, Hezel S, Simon JC. Topical treatment of atopic dermatitis with St. John's wort cream--a randomized, placebo controlled, double blind half-side comparison. *Phytomedicine.* 2003;10 Suppl 4:31-37. <https://doi.org/10.1078/1433-187x-00306>.
- [70] Hoffmann J, Wölflle U, Schempp CM, Casetti F. Tannins from *Potentilla officinalis* display antiinflammatory effects in the UV erythema test and on atopic skin. *J Dtsch Dermatol Ges.* 2016;14(9):917-922. <https://doi.org/10.1111/ddg.12792>.
- [71] Lin YK, Chang CJ, Chang YC, Wong WR, Chang SC, Pang JH. Clinical assessment of patients with recalcitrant psoriasis in a randomized, observer-blind, vehicle-controlled trial using indigo naturalis. *Arch Dermatol.* 2008;144(11):1457-1464. <https://doi.org/10.1001/archderm.144.11.1457>.
- [72] van de Kerkhof PC, van der Valk PG, Swinkels OQ, Kucharekova M, de Rie MA, de Vries HJ, Damstra R, Oranje AP, de Waard-van der Spek FB, van Neer P, Lijnen RL, Kunkeler AC, van Hees C, Haertlein NG, Hol CW. A comparison of twice-daily calcipotriol ointment with once-daily short-contact dithranol cream therapy: a randomized controlled trial of supervised treatment of psoriasis vulgaris in a day-care setting. *Br J Dermatol.* 2006;155(4):800-807. <https://doi.org/10.1111/j.1365-2133.2006.07393.x>.
- [73] Saller R, Büechli S, Meyrat R, Schmidhauser C. Combined herbal preparation for topical treatment of Herpes labialis. *Forsch Komplementarmed Klass Naturheilkd.* 2001;8(6):373-382. <https://doi.org/10.1159/000057255>.
- [74] Pflugfelder A, Andonov E, Weide B, Dirschka T, Schempp C, Stockfleth E, Stratigos A, Krüger-Krasagakis S, Bauer J, Garbe C, Eigentler TK. Lack of activity of betulin-based Oleogel-S10 in the treatment of actinic keratoses: a randomized, multicentre, placebo-controlled double-blind phase II trial. *Br J Dermatol.* 2015;172(4):926-932. <https://doi.org/10.1111/bjd.13342>.
- [75] Laszczyk M, Jäger S, Simon-Haarhaus B, Scheffler A, Schempp CM. Physical, chemical and pharmacological characterization of a new oleogel-forming triterpene extract from the outer bark of birch (*Betula cortex*). *Planta Med.* 2006;72(15):1389-1395. <https://doi.org/10.1055/s-2006-951723>.
- [76] Khazdair MR, Boskabady MH, Hosseini M, Rezaee R, M Tsatsakis A. The effects of *Crocus sativus* (saffron) and its constituents on nervous system: A review. *Avicenna J Phytomed.* 2015;5(5):376-391.
- [77] Devi S, Gupta A, Sahu M, Mishra P. Proven Health Benefits and Uses of Coriander (*Coriandrum sativum* L.) Series. In: *Ethnopharmacological Investigation of Indian Spices* Mishra N (editor). Kanpur, GI Global, 2020; 197-204
- [78] Kusindarta DL, Wihadmadyatami H, Haryanto A. *Ocimum sanctum* Linn. stimulate the expression of choline acetyltransferase on the human cerebral microvascular endothelial cells. *Vet World.* 2016; 9(12):1348-1354. <https://doi.org/10.14202/vetworld.2016.1348-1354>.
- [79] Hosseini M, Mohammadpour T, Karami R, Rajaei Z, Sadeghnia HR, Soukhtanloo M. Effects of the hydro-alcoholic extract of *Nigella sativa* on scopolamine-induced spatial memory impairment in rats and its possible mechanism. *Chin J Integr Med.* 2015; 21(6): 438-444. <https://doi.org/10.1007/s11655-014-1742-5>.
- [80] Bagheri SM, Rezvani ME, Vahidi AR, Esmaili M. Anticonvulsant effect of ferula assa-foetida oleo gum resin on chemical and amygdala-kindled rats. *N Am J Med Sci.* 2014;6(8):408-412. <https://doi.org/10.4103/1947-2714.139296>.
- [81] Pan HY, Qu Y, Zhang JK, Kang TG, Dou DQ. Antioxidant activity of ginseng cultivated under mountainous forest with different growing years. *J Ginseng Res.* 2013; 37(3):355-360. <https://doi.org/10.5142/jgr.2013.37.355>.
- [82] Paterna JC, Leng A, Weber E, Feldon J, Büeler H. DJ-1 and Parkin modulate dopamine-dependent behavior and inhibit MPTP-induced nigral dopamine neuron loss in mice. *Mol Ther.* 2007; 15(4):698-704. <https://doi.org/10.1038/sj.mt.6300067>.
- [83] Sbrini G, Brivio P, Fumagalli M, Giavarini F, Caruso D, Racagni G, Dell'Agli M, Sangiovanni E, Calabrese F. *Centella asiatica* L. Phytosome Improves Cognitive Performance by Promoting Bdnf Expression in Rat Prefrontal Cortex. *Nutrients.* 2020;12(2):355. <https://doi.org/10.3390/nu12020355>.
- [84] Eidi M, Eidi A, Bahar M. Effects of *Salvia officinalis* L.(sage) leaves on memory retention and its interaction with the cholinergic system in rats. *Nutrition.* 2006; 22(3):321-326. <https://doi.org/10.1016/j.nut.2005.06.010>.
- [85] Rahman K, Lowe GM. Garlic and cardiovascular disease: a critical review. *J Nutr.* 2006; 136: 736-740. <https://doi.org/10.1093/jn/136.3.736s>.
- [86] Satyavati GV, Dwarkanath C, Ttripathi SN. Experimental studies on the hypocholesterolemic effect of *Commiphora mukul* Engl. (Guggul). *Indian J Med Res.* 1969; 57(10): 1950-1962.
- [87] Chen JD, Wu YZ, Tao ZL, Chen ZM, Liu XP. Hawthorn (shanzha) drink and its lowering effect on blood lipid levels in humans and rats. *World Rev Nutr Diet* 1995; 77: 147-154. <https://doi.org/10.1159/000424470>.
- [88] Dwivedi S. *Terminalia arjuna* Wight & Arn. – A useful drug for cardiovascular disorders. *J Ethnopharmacol.* 2007;114(2):114-129. <https://doi.org/10.1016/j.jep.2007.08.003>.

- [89] Baumann G, Felix S, Sattelberger U, Klein G. Cardiovascular effects of forskolin (HL 362) in patients with idiopathic congestive cardiomyopathy – a comparative study with dobutamine and sodium nitroprusside. *J Cardiovasc Pharmacol.* 1990; 16:93-100. <https://doi.org/10.1097/00005344-199007000-00013>
- [90] Seifen E, Adams RJ, Riemer RK. Sanguinarine: a positive inotropic alkaloid which inhibits cardiac Na⁺,K⁺-ATPase. *Eur J Pharmacol.* 1979; 60:373-377. [https://doi.org/10.1016/0014-2999\(79\)90245-0](https://doi.org/10.1016/0014-2999(79)90245-0).
- [91] Shahrjerdi K. Cardioprotective effect of *Zanthoxylum americanum* mill. on isoproterenol-induced cardiotoxicity in albino wistar rats. *World J Pharm Res.* 2008; 7(5): 705-713.
- [92] Kang DG, Yun YG, Ryoo JH, Lee HS. Anti-hypertensive effect of water extract of danshen on renovascular hypertension through inhibition of the renin angiotensin system. *Am J Chin Med.* 2002; 30:87-93. <https://doi.org/10.1142/s0192415x02000107>.
- [93] Hirasa K., Takemasa, M. Spice science and technology. Marcel Dekker, New York. 1998; p. 220.
- [94] Dorman HJD, Deans SG. Antimicrobial agents from herbals: antibacterial activity of herbal volatile oils. *J Appl Microbiol.* 2000; 88: 308-316. <https://doi.org/10.1046/j.1365-2672.2000.00969.x>.
- [95] Newton SM, Lau C, Gurucha SS, Besra GS, Wright CW. The evaluation of forty-three plant species for in vitro antimycobacterial activities: Isolation of active constituents from *Psoralea corylifolia* and *Sanguinaria canadensis*. *J Ethnopharmacol.* 2002;79(1):57-67. [https://doi.org/10.1016/s0378-8741\(01\)00350-6](https://doi.org/10.1016/s0378-8741(01)00350-6).
- [96] Scalbert A. Antimicrobial properties of tannins. *Phytochemistry.*1991; 30: 3875-3883. [https://doi.org/10.1016/0031-9422\(91\)83426-L](https://doi.org/10.1016/0031-9422(91)83426-L).
- [97] Huyghebaert G., Ducatelle R, Van Immerseel F. 2011. An update on alternatives to antimicrobial growth promoters for broilers. *Vet J.* 2011;187(2):182-188. <https://doi.org/10.1016/j.tvjl.2010.03.003>.
- [98] Muanda F, Kone D, Dicko A, Soulimani R, Younos C. Phytochemical Composition and Antioxidant Capacity of Three Malian Medicinal Plant Parts. *Phytochemical composition and antioxidant capacity of three malian medicinal plant parts. Evid Based Complement Alternat Med.* 2011;2011:674320. <https://doi.org/10.1093/ecam/nep109>.
- [99] Liu HW, Dong XF, Tong JM, Zhang Q. Alfalfa polysaccharides improve the growth performance and antioxidant status of heat-stressed rabbits. *Livestock Sci.* 2010; 131: 88-93. <https://doi.org/10.1016/j.livsci.2010.03.004>.
- [100] Frankic T, Voljg M, Salobir J, Rezar V. Use of Herbs and spices and their extracts in animal nutrition. *Acta Agric Slov.* 2009; 92(2): 95-102.
- [101] Botha CJ, Penrith ML. Poisonous plants of veterinary and human importance in southern Africa. *J Ethnopharmacol.* 2008; 119:549-558. <https://doi.org/10.1016/j.jep.2008.07.022>.
- [102] Gaillard Y, Pepin G. Poisoning by plant material: review of human cases and analytical determination of main toxins by high-performance liquid chromatography-(tandem) mass spectrometry. *J Chromatogr B Biomed Sci Appl.* 1999;733(1-2):181-229. [https://doi.org/10.1016/s0378-4347\(99\)00181-4](https://doi.org/10.1016/s0378-4347(99)00181-4).
- [103] Štefanac T, Grgas D, Landeka Dragičević T. Xenobiotics-Division and Methods of Detection: A Review. *J Xenobiot.* 2021 Oct 26;11(4):130-141. doi: 10.3390/jox11040009. PMID: 34842778; PMCID: PMC8628977.
- [104] Nair B. Final report on the safety assessment of Mentha Piperita (Peppermint) Oil, Mentha Piperita (Peppermint) Leaf Extract, Mentha Piperita (Peppermint) Leaf, and Mentha Piperita (Peppermint) Leaf Water. *Int J Toxicol.* 2001;20 Suppl 3:61-73.
- [105] Sudekum M, Poppenga RH, Raju N, Braselton WE Jr. Pennyroyal oil toxicosis in a dog. *J Am Vet Med Assoc.* 1992;200(6):817-818.