# Nature's Cure or Kidney Curse? The Nephrotoxic Potential of Indigenous Remedies

# Abstract

Traditional medicine refers to a diverse range of health and healing practices used by Indigenous peoples that incorporate plants, animals, and the spiritual realm for the diagnosis and treatment of illness or maintenance of health and well-being. While these practices hold significant cultural value, there is growing evidence of potential nephrotoxicity associated with some traditional indigenous medicines (TIMs). In this review, we explore the various clinical manifestations of nephrotoxicity induced by TIMs. Nephrotoxicity can result from various factors, including the intrinsic toxicity of certain herbs, contamination with heavy metals, inappropriate dosages, and interaction with conventional medications. This review highlights the importance of monitoring kidney function in patients using TIMs and calls for increased awareness and regulation to ensure the safe use of these traditional therapies.

Keywords: Acute kidney injury, Herbal, Indigenous, Nephrotoxicity, Traditional medicines

### Introduction

Disparities in healthcare access in the developing world often lead to the use of traditional medicines in the treatment of not only common ailments but serious medical conditions as well. These traditional medicines have been associated with nephrotoxicity but rarely documented in modern literature, as these alternative medical systems and therapies are more often prevalent in the remote areas of developing countries.<sup>1,2</sup> Herbs are the most common constituent of these remedies and are often sourced locally and the medications prepared in-house.<sup>3</sup> The World Health Organization (WHO) estimates that 80% of the population of Asian and some African countries presently use herbal medicine for some aspect of primary healthcare. In developing countries like India, 65% of rural communities still use a traditional form of medicine to fulfill their primary healthcare needs.4 The WHO and the Government of India recently signed an agreement to establish the WHO Global Centre for Traditional Medicine that aims to harness the potential of traditional medicine from across the world through modern science and technology. The National Rural Health Mission has also decided to mainstream the Ayurveda, Yoga, Naturopathy, Unani, Siddha, and

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Homeopathy (AYUSH) system of indigenous medicine to help meet the challenge of shortage of healthcare professionals and strengthen the delivery system of the healthcare service in India.<sup>5</sup>

These traditional medicines are cheap, and considered accessible, natural, relatively safer than modern medicines. However, with the increasing use of these medications, the incidence of nephrotoxicity is becoming increasingly apparent. It is estimated that traditional remedies account for up to 35% of cases of acute kidney injury (AKI) in the developing world.<sup>2</sup> Even in developed countries, herbal medicines accounted for approximately 26% of all alternative medicine use.<sup>3</sup> The true incidence of chronic kidney disease (CKD) secondary to herbal medicines is not known because of the challenges in the diagnosis of CKD in this subset of patients.<sup>3</sup> In a survey from Africa, the prevalence of traditional medicine use among adults with CKD was 70.3%.<sup>6</sup> A cross-sectional study from Taiwan showed that herbal therapy use was associated independently with the development of CKD.7

Various epidemiological studies and chemical examination of herbal medicines have led to the identification of the culprit chemical compound, as illustrated in

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cases of AA-associated Chinese herb nephropathy/Balkan nephropathy.<sup>8</sup> Besides various organic acids and alkaloids, use of heavy metals (deliberately used for contamination during production) in traditional herbal medications have been implicated in nephrotoxicity.<sup>9</sup> Renal syndromes associated with traditional indigenous medicines (TIMs) include dyselectrolytemia, Fanconi syndrome, AKI, rhabdomyolysis, CKD, nephrolithiasis, urothelial malignancy, proteinuric kidney disease, and lupus nephritis. Figure 1 depicts the different clinical syndromes caused by indigenous medicines.



**Figure 1:** Various kidney syndromes caused by traditional medicines. AKI: Acute kidney injury, CKD: Chronic kidney disease, CIN: Chronic interstitial nephritis.

## Nephrotoxic Clinical Syndromes with Indigenous Medications

### **Electrolyte disturbances**

Electrolyte disturbances can occur in the use of herbal medications. Unauthorized herbal juices can cause hyperkalemia, especially in patients with CKD. Use of noni juice (*Morinda citrifolia*) has been shown to cause hyperkalemia, because the herb contains a very high level of potassium (56.3 meq/L) similar to that of orange or tomato juice.<sup>10</sup> Besides hyperkalemia, alternative medication use<sup>10–15</sup> has been shown to cause other forms of electrolyte abnormalities [Table 1].

### Fanconi syndrome

Fanconi syndrome was one of the earliest renal manifestations recognized with the use of *Aristocholic Acid* (AA), wherein the use of Chinese herbs leads to the clinical symptoms of hypokalemic periodic paralysis. A Chinese herbal remedy (Akebia 14) has been reported to cause reversible Fanconi syndrome. This medicine, marketed as herbal weight loss supplement, also contains high quantities of AA. Patients on these herbal remedies usually present with severe body pains, myopathy, and on evaluation have hypophosphatemia, hypouricemia, glucosuria, generalized aminoaciduria, and tubular proteinuria, the five cardinal symptoms of the Fanconi syndrome. Renal biopsy done in some of these reports has shown interstitial edema, focal tubular atrophy fibrosis, and glomerular ischemic damage. Usually, Fanconi syndrome is reversible when the herbal

# Table 1: Electrolyte abnormalities caused by traditional medicines

Alternative product/herb	Electrolye/acid base abnormality	Indication of use
Glycyrrhiza glabara (licorice)	Hypokalemia/ pseudoaldosteronism, hypertension, and metabolic alkalosis	Chinese herbal tea marketed as cough suppressant
		Korean herbal tonic marketed for prostate cancer
Cesium chloride	Hypokalemia	Alternative and complementary cancer treatment
<i>Morinda citrifolia</i> (noni juice)	Hyperkalemia	Nutritional supplement
Serenoa repens (saw palmetto)	Hyperkalemia	Remedy for benign prostatic hyperplasia
Oolong tea (detox tea)	Hyponatremia	Weight loss supplement
Colon cleanser	Hyponatremia	Enema

drug is withdrawn. In some cases, prolonged use of these herbs has led to progressive renal failure.<sup>16,17</sup>

### Rhabdomyolysis

Rhabdomyolysis, secondary to the intake of herbal medications, have been reported in literature, especially with medications prescribed for weight loss. Case reports of increasing creatinine phosphokinase (CPK) associated with herbal medicines are also discussed in literature. Elevated CPK is indicative of muscle breakdown, which is the mechanism by which these herbal medications achieve weight loss.<sup>18</sup> Many nonregulated herbal medications used for regulating menstrual cycles (Vitex) and alleviating some of the premenstrual symptoms have shown to cause rhabdomyolysis.<sup>19</sup> In other instances, herbal drugs interact adversely with modern medicines and can cause rhabdomyolysis. CPK should be checked in all patients with muscle aches and pains in patients on herbal medications. In most cases, the muscle destruction reverses with the discontinuation of the traditional medicine. Rhabdomyolysis, caused by contaminated/ substituted herbal hair dyes, can cause severe AKI and death. Physicians should be aware of the potential side effects of many herbal medicines and measure serum CPK enzyme for patients who admit using weight loss herbs and present with AKI.

#### Acute kidney injury

AKI is a common manifestation with the use of traditional medicines. The acute injury can manifest either as acute tubular necrosis or as acute interstitial nephritis. Indigenous medicines form the backbone for the treatment of common ailments like menstrual problems, fertility issues, malaria, typhoid, and more in developing nations. Treatment with these unregulated therapies often causes AKI. Around one-third cases of AKI are caused by the consumption of traditional medications.<sup>20,21</sup> Similar situation exists in rural India where a majority of the population depend on traditional healers and therapies.<sup>21</sup>

AKI may be the only manifestation of the herbal medications. In certain cases, AKI is accompanied with hepatotoxicity, neurological manifestations, bleeding diathesis, and respiratory failure. Patients usually present with severe gastroenteritis, dehydration, and shock. AKI is usually consequent to severe intravascular volume depletion. These patients often have very high mortality.<sup>22</sup> In other instances, AKI is secondary to the contamination of the traditional folk remedy with nephrotoxic chemicals. A classic example of AKI being secondary is the ingestion of "holy water." This liquid is often laced with copper sulphate giving its distinct color which leads to intravascular hemolysis and AKI.23 Contamination with potassium bromide, potassium dichromate, aloe, phyllum resin, phenolphthalein, and carbon tetrachloride has also been reported in literature.<sup>24</sup> Adulteration of traditional medicines happens during the process of manufacture. Many over-the-counter Chinese herbal and nutritional supplements are mixed with ephedrine, chlorpheniramine, sildenafil, steroids, fenfluramine, phenacetin, and so on and so forth. Ten percent of these supplements have undeclared drugs and heavy metals.<sup>25,26</sup> This chemical adulteration is done to enhance the efficacy of the remedy, and in some instances, the contamination is unintentional. This accidental contamination of herbal medicines happens when these plants are grown in highly polluted soil.<sup>27</sup>

AKI can also develop if the traditional remedies are introduced into the human body in an unusual manner. Use of herbal enemas in Africa have been associated with acute renal failure. Herbal enemas are very popular, and members of the Zulu tribe often have three enemas a week. These practices cause colonic complications leading to dehydration, infection, bleeding, and shock. AKI is a common consequence.<sup>28</sup>

Occasionally, AKI is seen when the original herbal product is replaced with a natural substitute. Mixing henna with paraphenylenediamine in hair dye can cause negative effects. Severe allergic reactions and rhabdomyolysis is reported leading to AKI.<sup>29</sup> In Morroco, hair is often dyed with a powder made from takaout el badia (made from the seeds of *Tamaris orientalis*). In situations of scarcity, the original powder is replaced with takaout roumia. Takaout roumia contains paraphenylenediamine which is responsible for rhabdomyolysis and AKI.<sup>30</sup> This product is responsible for 10% of AKI in Morrocco.

Another common situation where AKI develops is when the patient simultaneously consumes modern medicines and traditional herbs. The interaction between the drugs sometimes leads to unexpected renal consequences. Use of cyclosporine and St John's wort (derived from *Hypericum*  *perforatum*) causes induction of cyclosporine metabolism. St John's wort induces cytochrome P 450 enzyme, the enzyme responsible for cyclosporine metabolism. Low cyclosporine levels can precipitate allograft rejection and AKI.<sup>31</sup>

Artemisia herba (white wormwood) is used in folk medicine in many parts of Europe for its antihelmintic/ antirheumatic properties. It can lead to AKI by osmotic nephrosis. Similarly, extracts of Euphorpbia paralias (sea spurge) is used in Europe and North Africa for treating cancer and infections. This herb can lead to AKI due to severe acute tubular necrosis.<sup>32</sup>

Besides herbs, animal products and extracts have been also implicated in the development of AKI.<sup>33–40</sup> Common example is propolis, a resinous product sourced by honey bees to build their hives.This substance is used as an antiseptic anti-inflammatory agent, anesthetic, antioxidative, hepatoprotective substance in South America over the ages. Various chemicals like aromatic acids, esters, flavonoids, benzyl cinnamate, methyl cinnamate make up propolis, but it is difficult to pinpoint the guilty chemical compound responsible for AKI. Even though systemic toxicity secondary to use of propolis is rare, there are case reports of acute renal failure following propolis ingestion.<sup>41</sup>

Gall bladder extract of various fishes, especially carp, is commonly used in folk therapies in Asia (China, Taiwan, Hong Kong, Japan, South Korea, and India). The fish bile is used as a cough expectorant as well as for its antipyretic, antihypertensive and anti-allergic effects. Symptoms appear very early after ingestion of the offending bile, which includes watery diarrhea, jaundice, bradycardia ,hypotension, and oligoanuria.The development of AKI is attributed to the cardiotoxicity of bile. Renal failure sets in within two days and often lasts for two to three weeks.<sup>42</sup>

Bile extract of sheep is also consumed for its medicinal value in Saudi Arabia. It is used as a treatment for diabetes mellitus. Use of sheep bile daily over weeks causes nausea and diarrhea. Intravascular volume depletion is responsible for the acute decline in kidney function. AKI is often associated with hyponatremia and metabolic encephalopathy.<sup>43</sup> Cases declined after the intake of sheep bile was regulated by the government of Saudi Arabia.

## Chronic kidney disease

Acute deterioration of renal function secondary to folk medicine is easy to recognize the progressive decline of renal function because the long-term consumption is often difficult to detect. Hence, the incidence of CKD secondary to herbal medicines is underreported. Despite the limitations in establishing causality, there are several reports of herbs which have been implicated in the development of CKD.<sup>3</sup> The chronic decline can be because of parenchymal fibrosis or may be a manifestation of nephrolithiasis. Long-standing CKD can progress to the acquired cystic kidney disease or the offending medication can cause cysts *de novo*. An undesirable and unwanted consequence of herbal medicine-induced CKD is the development of urothelial carcinoma. The commonest and most well-researched herb leading to the development of CKD is AA, a toxic constituent of the Chinese herb nephropathy.

Besides AA, there are other herbs which have been associated with progressive irreversible deterioration of kidney function. Leaves of a creosote bush (Larrea tridentata) are commonly used as tea in North America, popularly known as chaparral tea. The roots and leaves are also marketed in the form of capsules and tablets. The active substance nordihydroguaiaretic acid is thought to have anticancer properties. Prolonged consumption of chaparral tea has led to the development of renal cysts and CKD.<sup>44,45</sup> In the modern world, licorice (*Glycyrrhiza* glabra) is a common constituent of lozenges and its extract is commonly used in alleviating the symptoms of epigastric discomfort and heartburn. It is known to cause hypokalemia (apparent mineralocorticoid excess). Prolonged hypokalemia leads to the development of hypokaemic nephropathy (a form of progressive interstitial injury).<sup>46</sup> Yohimbine, an extract from *Pausinystalia* yohimbe, is used as a folk remedy for erectile dysfunction. Use of this herbal extract causing a clinical illness similar to systemic lupus erythematosus and CKD has been reported in literature.<sup>47</sup> Bladderwrack is a large brown algae (Fucus vesiculosus) used as food in Japan. In Europe, it is marketed as herbal medicine for the treatment of atherosclerosis, heartburn, constipation, bronchitis, emphysema, urinary tract disorders, anxiety, and blood cleansing. CKD has been reported when bladderwrack tablets are consumed long term.<sup>48</sup> Similar manifestation of chronic decline of kidney function is noted when the Chinese herb rhubarb containing anthraquinone derivatives (Rhizoma rhei) is used as a slimming pill.<sup>49</sup>

The autopsy of the famous music composer Ludwig van Beethoven revealed the presence of papillary necrosis.<sup>50</sup> Papillary necrosis was postulated to be because of the chronic intake of willow bark (*Salix daphnoides*). The bark contains salicin which is metabolized to a well-known prostaglandin inhibitor—salicylate. Chronic ingestion of salicylic acid is known to cause papillary necrosis (chronic analgesic nephropathy).

Traditional herbal supplements may contain ethylene or diethylene glycol-based solvents that may lead to chronic oxalate nephropathy.<sup>51</sup> A thorough medication history should be obtained in patients with rapid decline of kidney function, even in the presence of known risk factors for CKD.

# Chronic kidney disease of unknown etiology and herbal medications

In the last decade, clusters of cases with unexplained CKD were reported in various parts of the world (Central

America, India, Sri Lanka, etc.).<sup>52,53</sup> These clusters were often in hot tropical climates, which predominantly affected farmers. These groups were in the underdeveloped part of the world with limited access to state-of-the-art healthcare facilities. The cause of CKD of unknown origin is still not clear and appears to be multifactorial. The World Health Organization realizes that the relatively common use of herbal medications in these affected areas may be contributing in the decline of renal function and has recommended restricting the use of herbal medicines in these regions.<sup>54</sup>

## **Glomerular disease**

Glomerular disease, secondary to the use of traditional Indian medications, has been reported in literature. The development of florid membranous nephropathy (MN) and minimal change disease (MCD), secondary to medicines prepared by practitioners of siddha and Ayurveda (a form of traditional medicine), has gained attention with the publication of reports in the last two years. These traditional healers combine herbs and inorganic chemical compounds such as sulfur and mercury in their medicinal preparations. The toxicity of these remedies is often because of the metal constituent. The drugs were used for a variety of indications like sinusitis, skin lesions, bronchitis, and breast cancer. These TIMs have been associated with neural epidermal growth factor-like 1 (NELL-1)-related membranous glomerulonephritis (MGN). Analysis of Swasa kalpa (a common Siddha medicine) showed higher than permissible mercury content.55 Previously, another report from India documented the development of MN in patients who were taking a herbo-mineral combination called rasagenthi mezhaku. The mercury content of this indigenous medication varied from 0.9% to 1.31%. The drug is available as 500 mg capsule and the prescribed dose is two tablets a day. The daily intake of two tablets a day led to a daily intake of 9-13.1 mg of mercury, which is much higher than the daily permissible limit (0.003 mg/day).<sup>56</sup> Heavy metals are mixed with herbs in various folk remedies not only in India, but in other parts of the world as well. In some studies, the levels of nephrotoxic heavy metals are more than the safe limits as required by regulatory authorities.<sup>57</sup>

#### Lupus nephritis

Indigenous medicines have immunomodulatory potential. Use of these drugs has been associated with various autoimmune conditions. CKD with a lupus-like presentation has been reported.<sup>47</sup> A recent report documents the development of lupus nephritis in a patient on herbal medicine. The patient was taking a drug called "super kidney" for diuresis. The unregistered drug contained ginseng, *Plantaginis folium*, orthosiphonis, *Strobilanthi folium*, and *Retrofacti fructus*. The patient's clinical condition did not completely improve with the cessation of the offending drug and she required immunosuppressive treatment.<sup>58</sup>

## Nephrolithiasis

Herbal medicines have been extensively researched for their antilithogenic potential. Some herbs when used inappropriately can cause both acute crystal-induced AKI and chronic nephrolithiasis. The most common example of crystal-induced tubular obstruction is secondary to the consumption of Djenkol beans-Archidendron pauciflorum (Djenkolism). Acute toxicity of Djenkol beans can cause hematuria and AKI.<sup>59</sup> Djenkolic acid is a sulfur-rich cysteine thioacetal of formaldehyde and forms intratubular crystals in the distal tubule of the nephron. These crystals then form the nidus for renal calculi (Djenkolic acid stones) formation. Progressive urolithiasis causes obstructive uropathy and chronic interstitial nephritis.<sup>60,61</sup> Another herbal medicine implicated in kidney stone formation is an appetizing fruit called star fruit (Averrhoa carambola). This fruit is used in the therapy of skin ailments and anxiety. Star fruit has very high levels of oxalic acid and can cause stones if consumed in large quantities.<sup>62</sup> The Chinese medicinal herb Ma Huang-derived Ephedra sinica is used for the treatment of bronchitis and nasal congestion. The chemical constituent Ephedrine is widely used in modern medicines. It is not only a popular medicine in China but also in India, and is considered to be the "somarasa" in Rigveda. Consumption of Ma Huang in large quantities with very little water can cause renal stone formation.63,64 Cranberry (Vaccinium macrocarpon) juice is widely considered a popular home remedy for recurrent urinary tract infection in the modern world. However, regular use of the concentrate which has high levels of oxalic acid can potentially cause urolithiasis.65

## **Urothelial carcinoma**

Urothelial malignancies happen when the transitional cell layer of the urinary tract mutate and multiply in an unregulated manner. Besides smoking, exposure to chemicals plays an important causative role. Environmental exposure of carcinogenic aromatic amines can happen when vulnerable individuals are exposed to various chemical industries. Other common and rarely appreciated mechanism is the contamination of the soil with these chemicals. Also in countries where herbal medication's use is in vogue, bladder cancer has been caused by the mutations secondary to AA intake. The highest incidence of urothelial cancers in the upper urinary tract is in Taiwan, where herbal medicines containing AA are used extensively.<sup>66</sup> The epidemiology of these cancers is remarkably similar to that noted in Balkan countries.67 This increase in the incidence of urothelial cancer in the last two decades in Taiwan is because of the replacement of original herbs with Aristolochia manchuriensis and Aristolochea fangchi.68 Urothelial carcinoma is often a late complication of AA intake and often manifests in patients who have developed end-stage kidney disease. Also, the continuous consumption of Chinese herbs by

CKD patients on dialysis increases their risk of developing urothelial carcinoma.<sup>69</sup> There are studies where urothelial cancer has developed relatively early in CKD patients who were not yet on dialysis.<sup>70</sup> Over the last two decades, governments are imposing ban on Chinese herbal products containing AA as their mutagenic potential is becoming clearer. This regulatory practice is impacting the incidence of upper urothelial cancer favorably.<sup>71</sup> The herbs and their constituents causing various clinical syndromes is summarized in Table 2.

Table 2: Common herbal plants which are implicated as
the cause of AKI/CKD and the indications of use

Herbal plant	Clinical manifestation	Indication of use
Acute kidney injury		
Aloe capensis (cape aloe)	Abdominal pain, hemorrhagic gastroenteritis/acute tubular necrosis in biopsies	Hypertension, arthritis
<i>Callilepis laureola</i> (Impila)	Abdominal pain, diarrhea, acute tubular necrosis, oliguric AKI, hepatic dysfunction, hypoglycemia, metabolic acidosis	To induce fertility, to correct impotence, and to ward off "evil spirits"
<i>Catha edulis</i> (Khat)	Acute tubular necrosis	Khat leaf chewing is a local habit in some African populations. Also used as folk remedy for obesity and depression
Colchicum	Gastroenteritis/	Used for acute
autumnale (Meadow saffron)	multiorgan failure/ hepatic injury/shock/ AKI	gouty arthritis- Suicidal colchicine tablet overdose is the most common cause of colchicine poisoning
Glycyrrhiza glabara (Licorice)	Contains glycyrrhetinic acid- can lead to hypokalemia, rhabdomyolysis, ARF, pseudoaldosteronism	Cough remedy
Amanita phalloides	Gastroenteritis,	Accidental ingestion,
(Toxic mushrooms)	hepatic failure, AKI	Anti-cancer, anti- inflammatory
<i>Caprica papaya</i> (papaya leave)	Severe mucositis and gastritis leading to severe dehydration causing pre-renal AKI	To increase platelet count in dengue fever
<i>Tamaris orientalis</i> (Takaout el badia)	Can lead to rhabdomyolysis and AKI	Henna for hair dyeing
Hypericum perforatum ( St. John's Wort)	Induces cytochrome P 450 enzyme-drug interactions and AKI	Used for depression and menopausal symptoms

Continued.,

## **Table 2: Continued**

Propolis (Honey bees product)	Acute interstitial nephritis	Used as antiseptic/ anti-inflammatory agent and hepatoprotective substance in South America
Carp (Fish gall bladder extract)	Can lead to watery diarrhea, jaundice, bradycardia, hypotension, oligoanuria and AKI	Used in folk therapies in parts of Asia
Caesalpinia decapetala (Cat's Claw)	Acute interstitial nephritis	A herbal remedy used in Peru as Anti- inflammatory drug for rheumatism, systemic lupus erythematosus, cirrhosis
<i>Cerbera thevetia</i> (Yellow Oleander)	Renal tubular necrosis	Used as folk remedy for heart conditions, asthma, epilepsy, cancer and painful menstrual periods
Cupressus funebris (Mourning Cypress)	Acute tubular necrosis, interstitial nephritis, and hemoglobin casts	Used as herbal remedy for bleeding piles and painful menstrual cycles
Securidaca longepedunculata (Wild Wisteria)	Contains salicylates-can cause ischemic renal damage/cortical necrosis and acute interstitial nephritis	Used as folk remedy for dysmenorrhea and venereal diseases
Euphorbia paralias (Spurge)	Tubular necrosis	Used as traditional medicine as an anti- inflammatory agent, a purgative and for its local anesthetic property.
Crotalaria laburnifolia (Bird flower) Obstructive AKI	Acute tubular injury	Used as Dysmenorrhea and Abortifacient
Archidendron pauciflorum (Djenkol beans)	Djenkolism-spasmodic suprapubic and/or flank pain, urinary obstruction and acute kidney injury	Local delicacy in Southeast Asia
Sobi-Lobi (Datura)	Contains anticholinergic substances-urinary obstruction and AKI	Consumed for its hallucinogenic properties
Salix daphnoides (Willow bark)	The bark contains salicin, which is metabolized in the body to salicylate-Renal papillary necrosis	Used for pain/ headache and inflammatory conditions in China and Europe

Continued.,

# Chronic kidney disease

**Table 2: Continued** 

Childrift Runey used	Se l	
Aristolochia clematitis (Birthwort)	Chronic interstitial nephritis, renal tubular defects, urothelial	Treatment of Gastrointestinal colic, slimming
Irvingia gabonensis (African Mango)	CKD-unknown mechanism	Used for weight reduction, diarrhea, dysentery
Lorrea tridentata (Chaparral)	Renal cystic disease-CKD	Herbal supplement marketed to treat colds, skin disorders, arthritis, bladder infections and cancer
Ephedra sinica (Ma-Huang)	Nephrolithiasis	Used for asthma, weight loss, sexual enhancement, and euphoria
Averrhoa Carambola (Star fruit)	Oxalate nephropathy/ Nephrolithiasis, obstructive nephropathy	Traditional remedy for diabetes mellitus in Asian countries
Vaccinium macrocarpon (Cranberry)	Oxalate nephropathy/ Nephrolithiasis, obstructive nephropathy	Used for reducing the risk of urinary tract infections
Pausinystalia yohimbe (Yohimbe)	Lupus nephritis	Used for erectile dysfunction, athletic performance and weight loss
Fucus vesiculosus (Bladder wrack)	Contains heavy metals- Chronic interstitial nephritis	Traditional herb used to treat reflux and heartburn
<i>Rhizoma Rhei</i> (Rhubarb)	Chronic interstitial nephritis	Traditional Chinese medication used to treat constipation/ cold sores
<i>Echinacea</i> spp. (Coneflower)	Renal tubular acidosis	Herbal remedy for common cold, cough, bronchitis and upper respiratory infections

AKI: Acute kidney injury, CKD: Chronic kidney disease, ARF: Acute renal failure

# Challenges in diagnosis of kidney disease due to traditional medicines

The prevalence of CKD has significantly increased in India over the past two decades. In a recent cross-sectional study from India, almost two-thirds of CKD patients were on some form of alternative medicine, in which Ayurveda was the most common.<sup>72</sup> Many individuals have a strong cultural or religious belief in the efficacy and safety of traditional medicines making them reluctant to disclose their use to healthcare providers. The clinicians mustactively seek information from their patients about the use of traditional/herbal drug remedies.

Undocumented traditional medicine ingestion should always be considered when investigating a patient with unexplained kidney disease. There is ample evidence to suggest that traditional medicines that were once considered natural and nontoxic contain some unknown toxic ingredients that might cause renal injury.<sup>73,74</sup> Public enlightenment is of utmost importance to educate the general population about the dangers posed by herbal remedies. Enforcing stringent regulations on the production and sale of traditional medicines can ensure better quality and safety standards.

As the popularity of herbal and traditional medicines continues to rise, the risks associated with their use cannot be overlooked. The concurrent use of traditional and modern medicines further complicates the landscape, posing challenges for diagnosis and treatment. Addressing this issue needs a multifaceted approach that includes enhancing public awareness about the potential risks, imposing strict regulatory frameworks, promoting better communication between patient and healthcare provider, and accessible/affordable healthcare services.

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