



Acta Universitaria

ISSN: 0188-6266

actauniversitaria@ugto.mx

Universidad de Guanajuato

México

Carranza-Álvarez, Candy; Alonso-Castro, Ángel Josabad; Maldonado-Miranda, Juan José; Hernández Morales, Alejandro

Quantitation of Cd, Pb and Fe in three medicinal plants (*Justicia spicigera*, *Arnica montana* and *Hamelia pantens*) from environmentally diverse locations of Huasteca Potosina, Mexico

Acta Universitaria, vol. 26, núm. 5, septiembre-octubre, 2016, pp. 24-30

Universidad de Guanajuato

Guanajuato, México

Available in: <http://www.redalyc.org/articulo.oa?id=41647917004>

- How to cite
- Complete issue
- More information about this article
- Journal's homepage in redalyc.org

redalyc.org

Scientific Information System

Network of Scientific Journals from Latin America, the Caribbean, Spain and Portugal

Non-profit academic project, developed under the open access initiative

Quantitation of Cd, Pb and Fe in three medicinal plants (*Justicia spicigera*, *Arnica montana* and *Hamelia pantens*) from environmentally diverse locations of Huasteca Potosina, Mexico

Cuantificación de Cd, Pb y Fe en tres plantas medicinales (*Justicia spicigera*, *Arnica montana* y *Hamelia pantens*) de distintas localidades de la Huasteca Potosina, México

Candy Carranza-Álvarez^{*♦}, Ángel Josabad Alonso-Castro^{**}, Juan José Maldonado-Miranda^{*}, Alejandro Hernández Morales^{*}

ABSTRACT

Recently, use of medicinal plants in Mexico and around the world has increased. However, in order to guarantee safe use of medicinal plants, it is necessary to quantify their content of heavy metals. In Mexico there is little information regarding content of toxic elements in medicinal plants, and these are consumed directly without a prior analysis of heavy metals. In this study, concentration of two toxic elements: cadmium and lead (Cd and Pb) and a micronutrient (iron [Fe]) was quantified in three medicinal plants (*Justicia spicigera*, *Arnica montana*, and *Hamelia pantens*) collected from three different locations in two municipalities in the Huasteca Potosina, Mexico. Cd and Pb are two toxic elements with no biological function, and can cause severe damage when introduced into food chain, whereas iron is an essential element. However, iron is a major element that can form oxides and entrap trace elements such as Pb and Cd, and induce bioaccumulation of these elements in plants. Also, in the Huasteca Potosina, the three medicinal plants of this study are used for empirical treatment of iron deficiency. Therefore, it was important to study concentration of the three elements in these medicinal plants. Results of this analysis showed that 50% and 22% of herbal preparations contain higher contents of Cd and Pb than those considered as safe, respectively. In addition, the three medicinal plants could be an important source of Fe. Results suggest that collection of medicinal plants should be carried out in sites free of potentially dangerous toxic elements.

RESUMEN

Recibido: 14 de septiembre de 2015
Aceptado: 17 de agosto de 2016

Keywords:

Medicinal plants; heavy metals; Cd; Pb; Fe.

Palabras clave:

Plantas medicinales; metales pesados; Cd; Pb; Fe.

Cómo citar:

Carranza-Álvarez, C., Alonso-Castro, Á. J., Maldonado-Miranda, J. J., & Hernández Morales, A. (2016). Quantitation of Cd, Pb and Fe in three medicinal plants (*Justicia spicigera*, *Arnica montana* and *Hamelia pantens*) from environmentally diverse locations of Huasteca Potosina, Mexico. *Acta Universitaria*, 26(5), 24-30. doi: 10.15174/au.2016.984

Recientemente, el uso de plantas medicinales en México y en todo el mundo se ha incrementado. Sin embargo, con el fin de garantizar el uso seguro en de las plantas medicinales, es necesario cuantificar su contenido de metales pesados. En México existe poca información sobre el contenido de elementos tóxicos en las plantas medicinales, y estas son consumidas directamente sin realizar un análisis previo de metales pesados. En este estudio se cuantificó la concentración de dos elementos tóxicos: cadmio y plomo (Cd y Pb) y de un micronutriente: fierro (Fe), en tres plantas medicinales (*Justicia spicigera*, *Arnica montana* y *Hamelia pantens*) colectadas de tres sitios diferentes de dos municipios de la Huasteca Potosina, México. El Cd y Pb son dos elementos tóxicos sin ninguna función biológica conocida, que pueden causar severos daños al introducirse en la cadena alimenticia, mientras que el hierro es un elemento esencial. Sin embargo, el hierro es un elemento mayoritario que puede formar óxidos y atrapar elementos traza como el Pb y Cd, y favorecer la bioacumulación de estos elementos en las plantas. Además, en la Huasteca Potosina, las tres plantas medicinales en estudio son usadas para el tratamiento empírico en la deficiencia de hierro. Por lo anterior, fue importante estudiar la concentración de los tres elementos en ambas plantas medicinales. Los resultados de este análisis mostraron que el 50% y el 22% de los preparados de plantas medicinales contienen valores más altos que los considerados como seguros para Cd y Pb, respectivamente. Además, se confirma que estas tres plantas medicinales podrían ser una fuente importante de Fe. Con esto se sugiere que la recolección de plantas medicinales se debe realizar en sitios libres de elementos tóxicos potencialmente peligrosos.

* Unidad Académica Multidisciplinaria Zona Huasteca, Universidad Autónoma de San Luis Potosí. Romualdo del Campo núm. 501, Fracc. Rafael Curiel, Ciudad Valles, S.L.P., México, C.P. 79060. Phone: (+52) 481 381 234 8; fax: (+52) 4813812348. E-mail: bqcca@hotmail.com; candy.carranza@uaslp.mx

** Departamento de Farmacia, División de Ciencias Naturales y Exactas, Campus Guanajuato, Universidad de Guanajuato, Guanajuato, México.

♦ Corresponding author.

INTRODUCTION

In the last two decades, use of herbal medicine has increased around the world, due to its putative efficiency, availability and general acceptability. Local flora is an important source of traditional medicines that are used for treatment of various diseases. For instance, 39 plant species belonging to 25 botanical families have been used for colon cancer treatment in Mexico with evidence in *in vitro* and *in vivo* studies (López-Rubalcava & Estrada-Camarena, 2016). In addition, 49 plants have been used in Mexican traditional medicine for treatment of disorders related to anxiety and depression (Jacobo-Herrera *et al.*, 2016), and more than 100 medicinal plants have been used as immunostimulants (Alonso-Castro, Juárez-Vázquez & Campos-Xolalpa, 2016). In urban areas, vendors of herbal products, called “yerbateros”, obtain medicinal plants from the wild and perform their prescription to general population. Therefore, in order to guarantee herbal products safe use, their content of toxic elements such as cadmium (Cd) and lead (Pb) should be analyzed. The uptake of heavy metals by plant occurs via roots and foliage (deposition and adsorption) (Madaan, Mudgal, Mishra, Srivastava & Singh, 2011). Factors that influence accumulation and concentrations of toxic elements in a plant include: atmospheric depositions, bioavailability, nature of soil and water (pH and organic matter concentration), and site of collection. These factors along with harvesting and processing conditions (*e.g.*, utensils containing Pb such as containers used for grinding) can affect heavy metals content in medicinal plants (Logan, Goins & Lindsay, 1997; Nwoko & Mgbeahuruike, 2011). The uptake of heavy metals by plants can increase certain toxic elements harmful potential that might enter into food chain. Accumulation of heavy metals in plant tissues disrupts many metabolic processes by alteration of oxidant-antioxidant balance and inhibition in activity and function of several enzymes (Rusyniak *et al.*, 2010).

In Mexican traditional medicine, *Justicia spicigera*, *Arnica montana* and *Hamelia patens* are used for empirical treatment of several diseases (Alonso-Castro *et al.*, 2012). A general description, as well as medicinal and economic importance of these plants are listed in table 1. In this study, these three medicinal plants were collected in the municipalities of Tamazunchale and Ciudad Valles, located in the *Huasteca Potosina* (Mexico) (figure 1), which is a region located in the coastal plain of the Gulf of Mexico. Some toxicological studies in terms of levels of heavy metals in water from rivers from the *Huasteca Potosina* have been carried out. For instance, Wong-Argüelles (2009) determined that levels of Cd and Pb in water from rivers of *Huasteca Potosina*,

including rivers from the municipalities of Ciudad Valles and Tamazunchale (figure 1), were below than those allowed by Mexican Legislation for drinking water (NOM-127-SSA1-1994). However, Mejía-Saavedra & Díaz-Barriga (2008) reported that As levels in a river from the municipality of Tamazunchale were higher than those allowed by Mexican Legislation for drinking water. Mejía-Saavedra & Díaz-Barriga (2008) also showed that Cd, Pb, Hg, and As levels in sediments from the river in Tamazunchale were higher than those recommended by the Canadian Legislation for the protection of wildlife (Canadian Council of Ministers of the Environment [CCEM], 2000).

General population perceives medicinal plants as safe and with no toxic effects because of their natural origin (Alonso-Castro *et al.*, 2012). Content of contaminants in medicinal plants should be analyzed and regulated by health systems worldwide to help quality assessment of these products. Poisonings associated with presence of toxic metals in medicinal plants have been reported in some countries (Dunbabin, Tallis, Popplewell & Lee, 1992; Kákósy, Hudák & Náray, 1996; Markowitz *et al.*, 1994; Sadhu *et al.*, 2015). Therefore, it is necessary to perform studies regarding content of heavy metals in medicinal plants used in Mexican traditional medicine. Nevertheless, in Mexico there is limited information in terms of toxic elements content in medicinal plants. The objective of this study was to collect three medicinal plants (*Justicia spicigera*, *Arnica montana* y *Hamelia patens*) from two municipalities in the *Huasteca Potosina* region and analyze their Pb, Cd and Fe content.

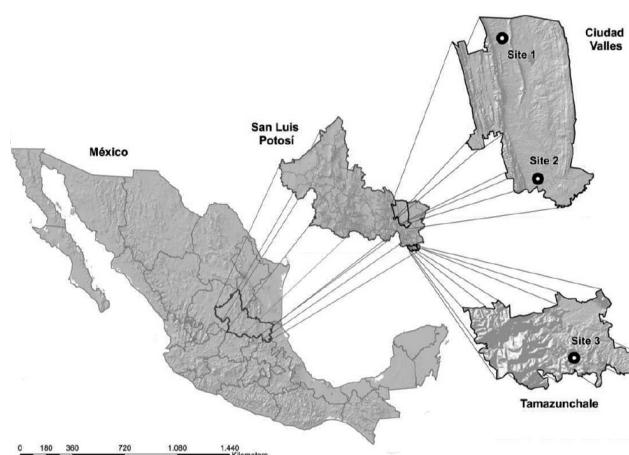


Figure 1. Map of study area, municipalities of Ciudad Valles and Tamazunchale, San Luis Potosí, México. Locations of the three sampling sites are shown. Source: Figure prepared by Geog. Enrique Ibarra Zapata.

Table 1.
General description and ethnopharmacological importance of medicinal plants used in the study.

Family	Scientific name voucher	Common name [Teenek name]	Part of plant used	Preparation/ Application	Therapeutic properties/targets	References
Rubiaceae	<i>Hamelia patens</i> Jacq. HPHP-SP-102	Chacloco [k'entselte']	Leaves	Infusion/oral	Antidiarrheal, intestinal antispasmodic, anemia, wound, healing, antidiabetic	Alonso-Castro <i>et al.</i> (2012), Cruz & Andrade-Ceto (2015).
Acanthaceae	<i>Justicia spicigera</i> Schltdl. JSHP- SP-43 15 1.19 Alliaceae Allium cepa L. ACHP- SP-81	Mohuite [Muu]	Leaves	Infusion/oral	Antidiabetic, prevents anemia, regulates high pressure, insomnia, wound healing, to keep away from bad spirits, emotional disorders	Alonso-Castro <i>et al.</i> (2012); Cassani <i>et al.</i> (2014); Ortiz-Andrade <i>et al.</i> (2012).
Asteraceae	<i>Arnica montana</i> L. ARHP-SP-47	Arnica [maan huitz]	Leaves	Infusion/oral	Antidiabetic, Antiprotozoal Activity antioxidant, anti-inflammatory, capacity body pain, rash, wounds, cough, arthritis, intestinal antispasmodic	Alonso-Castro <i>et al.</i> (2012); Llurba-Montesino, Kaiser, Brun & Schmidt (2015); Rodríguez-Chávez, Coballase- Urrutia, Nieto-Camacho & Delgado-Lamas (2015); Sharma, Arif, Nirala, Gupta & Thakur (2015).

Source: Author's own elaboration.

Table 2.
List of diverse locations in Huasteca Potosina, Mexico from where medicinal plants were collected.

Site	Municipality	District	Latitude	Longitude	Meters above sea level (m.a.s.l.)	Environmental characteristics of site
1	North side of of Ciudad Valles	San Luis Potosí	99° 01' W	21° 59' N	74	Cd and Pb in water from rivers
2	South side of Ciudad Valles	San Luis Potosí	99° 01' W	21° 50' N	70	Cd and Pb in water from rivers
3	Tamazunchale	San Luis Potosí	98° 48' W	21° 16' N	140	As, Hg, Cd and Pb in water from rivers

Source: Author's own elaboration.

MATERIALS AND METHODS

Whole plants of *J. spicigera*, *A. montana* and *H. patens* ($n = 3$), with a height of approximately 0.9 m to 1.0 m, were collected at three different sites in the Huasteca Potosina (table 2 and figure 2). The three plants were washed with tap water and then with deionized water to remove any adhering dust particles.

Plant material was sectioned into stem and leaves, dried at 70 °C during 1 h, and grounded in a Wiley mill for 2 min. Additionally, infusions of *J. spicigera*, *H. patens* and *A. montana* were prepared, each, by adding 1 g of aerial parts into 1 l of boiling distilled water. Plant material was boiled for 5 min and cooled to room temperature for 10 min. This procedure was carried

out according to traditional way infusion preparation of medicinal plants in the *Huasteca Potosina* (Alonso-Castro *et al.*, 2012). All samples were preserved in High-density polyethylene (HPDE) bottles at room temperature and protected from light and dust. Analyses of Cd, Pb and Fe were carried out by acid digestion with 0.1 mL HNO₃ (reactive grade, Fermont) per 2 mg of plant material during 4 days. Acid digestion was performed at room temperature in closed Teflon vessels. In order to complete organic matter oxidation, acid digestion was continued during 24 h by adding 0.5 mL of H₂O₂ 30% (v/v) (Alfaro-De La Torre, 1997; Yan, Mackie & Dillon, 1990). After evaporating remaining acid, samples were allowed to dry at room temperature, and residue was re-dissolved with HNO₃ 5% (v/v) to complete 25 mL (Loring & Rantala 1992).

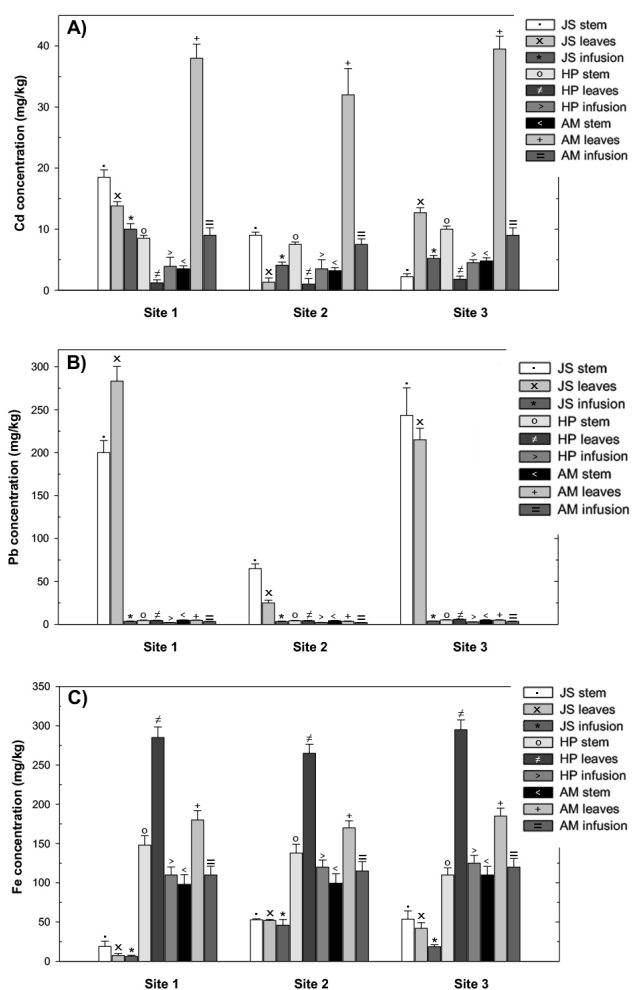


Figure 2. Mean ($n=3$) concentrations of heavy metals in leaves, stem and infusion of medicinal plants *Justicia spicigera*, *Hamelia patens* and *Arnica montana* collected from three different sites of Huasteca Potosina, México. A) Cd concentration (mg/kg); B) Pb concentration (mg/kg); C) Fe concentration (mg/kg). Error bars represent standard error of concentrations.

Source: Author's own elaboration.

Reference plant material *Lagarosiphon major* (Community Bureau of Reference [CBR], Reference material No. 60) and blanks were analyzed in triplicate to determine accuracy of determinations. Recovery rate for reference plant sample ranged between 90% and 110%.

Content of Cd, Pb, and Fe in plant material was determined using an atomic absorption spectrophotometer (AAS) using an air-acetylene flame or graphite furnace (Varian-SpectrAA 220 FS, Palo Alto, CA). Detection limits for analytical methods were: 0.01 mg/Kg, 0.01 mg/Kg, 0.03 mg/Kg, for Cd, Pb and Fe respectively. A certified water sample for trace elements

(TM-DWS, National Water Research, and Canada) was run with samples analyzed by AAS employing a graphite furnace to ensure an optimum quality control during analytical procedures. Recovery rate for certified water sample ranged between 100% \pm 15%.

Mean of metal concentrations in plant material is reported in mg/Kg based on dry weight. Differences between sampling sites with respect to mean concentrations of heavy metals in stem and/or leaves were evaluated by using a one-way analysis of variance (ANOVA) test at significance level of 95%. Results were considered significantly different where the calculated P-values were \leq 0.05. All calculations were done employing the Graph Instat TM V2.02 program.

RESULTS

Concentrations of Cd, Pb and Fe in raw material and their infusions are shown in figure 2. Findings indicated that the highest Cd concentration was found in leaves from *A. montana* collected in sites 1 (38 mg/kg \pm 2.3 mg/kg) and 3 (39.5 mg/kg \pm 2.1 mg/kg), whereas the lowest Cd concentration was found in *H. patens* leaves collected in site 2 (0.2 mg/kg \pm 0.01 mg/kg) (figure 2A). Safe limit for Cd is 0.3 mg/kg in plant material final dosage (World Health Organization [WHO], 1999). Taking this into account, the following preparations can be considered as safe: leaves and infusion of *H. patens* and *A. montana*, each from the 3 sites of collection, *J. spicigera* infusion from sites 2 and 3, *J. spicigera* leaves from site 2, and *J. spicigera* stem from site 3. In contrast, concentration of Pb in all medicinal plants ranged from 283.3 mg/kg (*J. spicigera* leaves from site 1) to 2.1 mg/kg (*A. montana* infusion from site 2) (figure 2B). Permissible limit for Pb is 10 mg/kg set by the WHO (1999). Considering this value, only *J. spicigera* leaves and stem, from all 3 sites of collection could not be considered as safe for human because it exceeds permissible limits of heavy metals in herbal products (Food and Agriculture Organization [FAO] / World Health Organization [WHO], 1984; WHO, 1998, 2004, 2007, 2011; Zhang 1998). Pb and Cd are toxic pollutants that affect human health (Khan *et al.*, 2014). Pb can cause anemia, neurological disorders, hyperactivity, among others (Marsden, 2003), whereas Cd affects lungs, reproductive system, kidneys, and bones (Godt *et al.*, 2006). In this study, concentrations of Pb in *J. spicigera* were higher than those found in medicinal plants from Ethiopia and Pakistan (Baye & Hymete, 2010; Nawab *et al.*, 2015). In addition, concentrations of Pb and Cd in the three medicinal plants were higher than those reported by Sadhu *et al.* (2015) in medicinal plants collected from

India. Iron (Fe) is an essential element for plants and humans. However, a chronic excessive ingestion of Fe induces hypovolemic shock and liver failure (Lee & Lee, 2002; Mudgal, Madaan, Mudgal, Singh & Mishra, 2010; Yi & Guerinet, 1996). Concentrations of Fe found in this study ranged from 6.3 mg/kg (*J. spicigera* infusion at site 1) to 295 mg/kg (*H. patens* leaves at site 3) (figure 2C). It is considered that content of Fe in plant tissue ranges from 50 mg/kg – 300 mg/kg (in terms of dry weight). Results suggest that these three medicinal plants contain an adequate concentration of Fe, which corroborates the empirical use of *J. spicigera* and *H. patens*, each, for empirical treatment of anemia (Alonso-Castro *et al.*, 2012). Nevertheless, it has been reported that plants exposed to Cd contamination decreases their uptake of Fe (Baryla *et al.*, 2001; Sikka & Nayyar, 2012). In this study, content of Cd found in medicinal plants did not alter their concentration of Fe.

It is probable that consuming herbal drugs obtained from polluted sources will lead to serious health hazards, especially if this results in exposure to Cd and Pb (Mishra, Dwivedi & Singh, 2010).

DISCUSSION

Heavy metals are natural occurring elements in the biosphere. Because of anthropogenic activities, many of these elements have been released in the environment and their concentrations have increased (Alloway, 1995). Worldwide, many areas used for agricultural purposes are contaminated with heavy metals (Nagajyoti, Lee & Sreekanth, 2010; Neilson & Rajakaruna, 2015). Plants can uptake toxic metals from the soil, water or air. Heavy metals may enter into food chain through ruminants, such as cattle, sheep, and goats, which are fed with plants (Oskarsson, Jorhem, Sundberg, Nilsson & Albanus, 1992; Waqas *et al.*, 2015). Human beings might be intoxicated with heavy metals by direct consumption of medicinal plants contaminated with these elements. Therefore, it is necessary to estimate presence of toxic elements such as Cd and Pb in medicinal plants in order to provide their safe use. Heavy metal concentrations were determined in raw material (stem and leaves) and in infusions of aerial parts of *J. spicigera*, *A. montana* and *H. patens* because these are plant parts and a way of administration commonly used in Mexican traditional medicine.

Accumulation of heavy metals in the three medicinal plants tend to vary depending on site of collection and on the medicinal plant. The highest concentrations of Cd were found in site A, whereas the high-

est concentrations of Pb were found in sites A and B. Some plants have developed mechanisms to grow in contaminated sites with heavy metals such as Cd and Pb (Khan *et al.*, 2015). In Fe case, there was not found a clear pattern of maximum accumulation among studied sites. Results also indicated the lowest accumulation of heavy metals in medicinal plants was found in site 2. Variations regarding accumulation of heavy metals in medicinal plants may be due to some factors such as mineral composition of soil, total concentration of dissolved heavy metals in each sampling site, predominant dissolved chemical species, and mechanisms of mobilization at sediment-water interface of heavy metals. Also, transfer coefficients of metals affect heavy metal accumulation in plants (Sadhu *et al.*, 2015). For instance, Pb has low transfer coefficients, remains stably bound to soil structures, and shows minimum bioavailability to plants. In contrast, metals like Cd have higher transfer coefficients and are easily taken up by plants (Kloke, Sauerbeck & Vetter, 1984).

Parameters described above are also influenced by release of chelating agents by plants as well as physicochemical and biological variations such as temperature, cation exchange capacity, nature of contaminants, salinity, sediment depth, redox potential, water and sediment pH, organic matter content, microbial biota, and physiological characteristics of each medicinal plant (Gregor, 2004).

CONCLUSION

In summary, this study indicates that detection of toxic elements, such as Cd and Pb, in medicinal plants is highly relevant for their quality control assessment. Therefore, medicinal plants should be monitored regarding place where they are collected. Also, it is necessary to implement policies in order to identify in marketplaces, herbs which contain contaminants, such as toxic metals, which represent a threat for human health. Thus, collection of medicinal plants should be done in monitored sites which are free of contamination. This study also gives information for conducting other studies in different parts of Mexico for quantification of toxic metals in medicinal plants.

ACKNOWLEDGEMENTS

This work was carried out with financial support from Apoyo a nuevos Profesores de Tiempo Completo del Programa de Mejoramiento de Profesorado (Prodep, Mexico) Grant No. 103.5/12/3953. We also thank to Dra. Maria Catalina Alfaro de la Torre (*Facultad de*

Química, Universidad Autónoma de San Luis Potosí, Mexico) for letting us to use her research facilities and the AAS equipment.

CONFLICT OF INTEREST

The authors declare that they have no conflict of interest.

REFERENCES

- Alfaro-De la Torre, M. C. (1997). Conteu des me'taux Cu, Pb, Cd et Zn dans les larves de L'insecte aquatique *Chaoborus* (Diptera) (30 pp.). *Rapport interne INRS-Eau Université du Québec, CAN*.
- Alloway, B. J. (Ed.) (1995). Soil processes and the behavior of metals. In *Heavy metals in soils* (pp. 38-57). London: Springer.
- Alonso-Castro, A. J., Juárez-Vázquez, M. C., & Campos-Xolalpa, N. (2016). Medicinal Plants from Mexico, Central America, and the Caribbean Used as Immunostimulants. *Evidence-Based Complementary and Alternative Medicine*, 2016, 1-15. doi: 10.1155/2016/4017676
- Alonso-Castro, A. J., Maldonado-Miranda, J. J., Zárate-Martínez, A., Jacobo-Salcedo, M. R., Fernández-Galicia, C., Figueroa-Zúñiga, L. A., Ríos-Reyes, N. A., De León-Rubio, M., Medellín-Castillo, N. A., Reyes-Munguía, A., Méndez-Martínez, R., & Carranza-Álvarez, C. (2012). Medicinal plants used in the Huasteca Potosina, Mexico. *Journal Ethnopharmacology*, 143(1), 292-298. doi: 10.1016/j.jep.2012.06.035
- Baryl, A., Carrier, P., Franck, F., Coulomb, C., Sahut, C., & Havaux, M. (2001). Leaf chlorosis in oilseed rape plants (*Brassica napus*) grown on cadmium-polluted soil: Causes and consequences for photosynthesis and growth. *Journal Planta*, 212(5-6), 696-709. PMID: 11346943.
- Baye, H., & Hymete, A. (2010). Lead and cadmium accumulation in medicinal plants collected from environmentally different sites. *Bulletin of Environmental Contamination and Toxicology*, 84(2), 197-201. doi: 10.1007/s00128-009-9916-0
- Canadian Council of Ministers of the Environment (CCME) (2000). Canadian tissue residue guidelines for the protection of wildlife consumers of aquatic biota: Methylmercury. In *Canadian environmental quality guidelines*, 1999. Winnipeg: CCME.
- Cassani, J., Dorantes-Barrón, A. M., Mayagoitia-Novales, L., Alva-Real, G. A., & Estrada-Reyes, R. (2014). Anti-depressant-like effect of kaempferitrin isolated from *Justicia spicigera* Schldl (Acanthaceae) in two behavior models in mice: evidence for the involvement of the serotonergic system. *Molecules*, 19(12), 21442-21461. doi: 10.3390/molecules191221442
- Cruz, E. C., & Andrade-Cetto, A. J. (2015). Ethnopharmacological field study of the plants used to treat type 2 diabetes among the Cakchiquels in Guatemala. *Journal of Ethnopharmacology*, 159, 238-244. doi: 10.1016/j.jep.2014.11.021
- Dunbabin, D. W., Tallis, G. A., Popplewell, P. Y., & Lee, R. A. (1992). Lead poisoning from Indian herbal medicine (Ayurveda). *Medical Journal of Australia*, 157(11-12), 835-836. PMID: 1454025
- Food and Agriculture Organization (FAO) / World Health Organization (WHO) (1984). Contaminants. In *Codex Alimentarius*, vol. XVII. Rome: FAO / WHO.
- Godt, J., Scheidig, F., Grosse-Siestrup, C., Esche, V., Brandenburg, P., Reich, A., & Groneberg, D. A. (2006). The toxicity of cadmium resulting hazards for human health. *Journal of Occupational Medicine and Toxicology*, 1(22), 1-6. doi: 10.1186/1745-6673-1-22
- Gregor, M. (2004). Metal availability, uptake, transport and accumulation in plants. In M.N.V. Prasad (Ed.), *Heavy metal stress in plants from biomolecules to ecosystems* (pp. 1-27). Berlin: Springer-verlag.
- Jacobo-Herrera, N. J., Jacobo-Herrera, F. E., Zentella-Dehesa, A., Andrade-Cetto, A., Heinrich, M., & Pérez-Plasencia, C. (2016). Medicinal plants used in Mexican traditional medicine for the treatment of colorectal cancer. *Journal of Ethnopharmacology*, 179, 391-402. doi: 10.1016/j.jep.2015.12.042
- Kákósy, T., Hudák, A., & Náray, M. (1996). Lead intoxication epidemic caused by ingestion of contaminated ground paprika. *Journal of Toxicology: Clinical Toxicology*, 34(5), 507-511. PMID: 8800188.
- Khan, K., Khan, H., Lu, Y., Ihsanullah I., Nawab, J., Khan, S., Shahe, N. S., Shamshad, I., & Maryama, A. (2014). Evaluation of toxicological risk of foodstuffs contaminated with heavy metals in Swat, Pakistan. *Ecotoxicology and Environmental Safety*, 108, 224-232. doi: 10.1016/j.ecoenv.2014.05.014
- Khan, S., Waqas, M., Ding, F., Shamshad, I., Arp, H. P. H., & Li, G. (2015). The influence of various biochars on the bioaccessibility and bioaccumulation of PAHs and potentially toxic elements to turnips (*Brassica rapa* L.). *Journal of Hazardous Materials*, 300, 243-253. doi: 10.1016/j.jhazmat.2015.06.050
- Kloke, A., Sauerbeck, D., & Vetter, H. (1984). The Contamination of Plants and Soils with Heavy Metals and the Transport of Metals in Terrestrial Food Chains. In J. O. Nriagu (Ed.), *Changing metal cycles and human health* (pp. 113-141). Berlin / Heidelberg: Springer. doi: 10.1007/978-3-642-69314-4_7
- Lee, S. M., & Lee, W. Y. (2002). Determination of heavy metal ions using conductometric biosensor based on Sol-gel immobilized urease. *Bulletin of the Korean Chemical Society*, 23(8), 1169-1172. doi: 10.5012/bkcs.2002.23.8.1169
- Llurba-Montesino, N., Kaiser, M., Brun, R., & Schmidt, T. J. (2015). Search for Antiprotozoal Activity in Herbal Medicinal Preparations; New Natural Leads against Neglected Tropical Diseases. *Molecules*, 20(8), 14118-14138. doi: 10.3390/molecules200814118
- Logan, T. J., Goins, L. E., & Lindsay, B. J. (1997). Field assessment of trace element uptake by six vegetables from N-Viro soil. *Water Environment Research*, 69(1), 28-33. doi: 0.2175/106143097X125155
- López-Rubalcava, C., & Estrada-Camarena, E. (2016). Mexican medicinal plants with anxiolytic or antidepressant activity: Focus on preclinical research. *Journal of Ethnopharmacology*, 186, 377-391.
- Loring, D. H., & Rantala, R. T. T. (1992). Manual for the geochemical analyses of marine sediments and suspended particulate matter. *Earth-Science Reviews*, 32(4), 235-283.
- Madaan, N., Mudgal, V. Mishra, S., Srivastava, A. K., & Singh, R. B. (2011). Studies on biochemical role of accumulation of heavy metals in Safflower. *The Open Nutraceuticals Journal*, 4, 199-204.

- Markowitz, S. B., Nenez, C. M., Klitzman, S., Munshi, A. A., Kim, W. S., Eisinger, J., & Landrigan, P. J. (1994). Lead poisoning due to haigefen: the porphyrin content of individual erythrocytes. *The Journal of the American Medical Association*, 271(12), 932-934. PMID: 8120963.
- Marsden, P. A. (2003). Increased body lead burden cause or consequence of chronic renal insufficiency? *New England Journal of Medicine*, 348(4), 345-347. PMID: 12540649.
- Mejía-Saavedra, J., & Díaz-Barriga, F. (2008). *Reporte interno*. Departamento de Toxicología Ambiental, Facultad de Medicina, Universidad Autónoma de San Luis Potosí, México.
- Mishra, S., Dwivedi, S. P., & Singh, R. B. (2010). A review on epigenetic effect of heavy metal carcinogens on human health. *The Open Nutraceuticals Journal*, 3, 188-193. doi: 10.2174/1876396001003010188. Retrieved September 9th 2015 from <http://www.benthamscience.com/open/tonutraj/articles/V003/188TONUTRAJ.pdf>
- Mudgal, V., Madaan, N., Mudgal, A., Singh, R. B., & Mishra, S. (2010). Effect of toxic metals on human health. *The Open Nutraceuticals Journal*, 3, 94-99. Retrieved September 10th 2015 from <http://www.benthamscience.com/open/tonutraj/articles/V003/94TONUTRAJ.pdf>
- Nagajyoti, P., Lee, K., & Sreekanth, T. (2010). Heavy metals, occurrence and toxicity for plants: A review. *Environmental Chemistry Letters*, 8(3), 199-216. doi: 10.1007/s10311-010-0297-8
- Nawab, J., Khan, S., Shah, M. T., Qamar, Z., Din, I., Mahmood, Q., Gul, N., & Huang, Q. (2015). Contamination of soil, medicinal, and fodder plants with lead and cadmium present in mine-affected areas, Northern Pakistan. *Environmental Monitoring and Assessment*, 187(9), 605-619. doi: 10.1007/s10661-015-4807-9
- Neilson, S., & Rajakaruna, N. (2015). Phytoremediation of agricultural soils: Using plants to clean metal-contaminated arable land. In A. A. Ansari, S. S. Gill, R. Gill, G. R. Lanza & L. Newman (Eds.), *Phytoremediation* (pp. 159-168). Cham, Switzerland: Springer International Publishing.
- Norma Oficial Mexicana (1994). NOM-127-SSA1-1994, *Salud ambiental, agua para uso y consumo humano. Límites permisibles de calidad y tratamientos a que debe someterse el agua para su potabilización*. México: Secretaría de Salud.
- Nwoko, C. O., & Mgbauruike, L. (2011). Heavy metal contamination of ready-to-use herbal remedies in south eastern Nigeria. *Pakistan Journal of Nutrition*, 10(10), 959-964.
- Ortiz-Andrade, R., Cabañas-Wuan, A., Arana-Argáez, V. E., Alonso-Castro, A. J., Zapata-Bustos, R., Salazar-Olivo, L. A., Domínguez, F., Chávez, M., Carranza-Álvarez, C., & García-Carrancá, A. (2012). Antidiabetic effects of *Justicia spicigera* Schldl (Acanthaceae). *Journal of Ethnopharmacology*, 143(2), 455-462. doi: 10.1016/j.jep.2012.06.043
- Oskarsson, A., Jorhem, L., Sundberg, J., Nilsson, N. G., & Abanus, L. (1992). Lead poisoning in cattle-transfer of lead to milk. *Science of the Total Environment*, 111(2-3), 83-94.
- Rodríguez-Chávez, J. L., Coballase-Urrutia, E., Nieto-Camacho, A., & Delgado-Lamas, G. (2015). Antioxidant capacity of "Mexican arnica" *Heterotheca inuloides* Cass natural products and some derivatives: Their anti-inflammatory evaluation and effect on *C. elegans* life span. *Oxidative Medicine of Cellular Longevity*, 1-11. doi: 10.1155/2015/843237
- Rusyniak, D. E., Arroyo, A., Acciani, J., Froberg, B., Kao, L., & Furbee, B. (2010). Heavy metal poisoning: management of intoxication and antidotes. In A. Luch (Ed.), *Molecular, Clinical and Environmental Toxicology* (pp. 365-396). Boston: Birkhäuser Basel. doi: 10.1007/978-3-7643-8338-1_11
- Sadhu, A., Upadhyay P., Singh, P. K., Agrawal, A., Ilango, K., Karmakar, D., Inder-Singh, G. P., & Prasad-Dubey, G. (2015). Quantitative analysis of heavy metals in medicinal plants collected from environmentally diverse locations in India for use in a novel phytopharmaceutical product. *Environmental Monitoring Assessment*, 187(8), 1-11. doi: 10.1007/s10661-015-4764-3
- Sharma, S., Arif, M., Nirala, R. K., Gupta, R., & Thakur, S. C., (2015). Cumulative therapeutic effects of phytochemicals in Arnica montana flower extract alleviated collagen-induced arthritis: inhibition of both pro-inflammatory mediators and oxidative stress. *Journal of the Science of Food and Agriculture*, 96(5), 1500-1510. doi: 10.1002/jsfa.7252
- Sikka, R., & Nayyar, V. (2012). Cadmium accumulation and its effects on uptake of micronutrients in Indian mustard [*Brassica juncea* (L.) Czern.] grown in a loamy sand soil artificially contaminated with cadmium. *Communications in Soil Science and Plant Analysis*, 43(4), 672-688. doi: 10.1080/00103624.2012.644007
- Waqas, M., Li, G., Khan, S., Shamshad, I., Reid, B. J., Qamar, Z., & Chao, C. (2015). Application of sewage sludge and sewage sludge biochar to reduce polycyclic aromatic hydrocarbons (PAH) and potentially toxic elements (PTE) accumulation in tomato. *Environmental Science and Pollution Research*, 22(16), 12114-12123. doi: 7071-7081. 10.1007/s11356-015-4432-8
- World Health Organization (WHO) (1998). *Quality control methods for medicinal plant materials*. Geneva Switzerland: WHO.
- World Health Organization (WHO) (1999). *Monographs on Selected Medicinal Plants*, vol. 1. Geneva, Switzerland: WHO.
- World Health Organization (WHO) (2004). WHO guidelines on safety monitoring of herbal medicines in pharmacovigilance systems. *Bulletin of the World Health Organization* (82 pp.) Geneva Switzerland: WHO. Retrieved September 14th 2015 from <http://apps.who.int/medicinedocs/documents/s7148e/s7148e.pdf>
- World Health Organization (WHO) (2007). *WHO guidelines for assessing quality of herbal medicines with reference to contaminants and residues*. Geneva, Switzerland: WHO. doi: ISBN 978 92 4 159444 8
- World Health Organization (WHO) (2011). *Quality Control Methods for Medicinal Plant Materials, Revised*. Geneva, Switzerland: WHO. doi: ISBN 978 92 4 150073 9
- Wong-Argüelles, C. (2009). *Estudio de organismos acuáticos macrobentónicos como indicadores de la contaminación por metales pesados en ríos de la Huasteca Potosina* (master of thesis). Programa Multidisciplinario de Posgrado en Ciencias Ambientales, Universidad Autónoma de San Luis Potosí: San Luis Potosí, México.
- Yan, N. D., Mackie, G. L., & Dillon, P. J. (1990). Cadmium concentrations at crustacean zooplankton of acidified and non acidified Canadian Shield lakes. *Environmental Science & Technology*, 24(9), 1367-1377.
- Yi, Y., & Guerinot, M. L. (1996). Genetic evidence that induction of root Fe (III) chelate reductase activity is necessary for iron uptake under iron deficiency. *The Plant Journal*, 10(5), 835-844. doi: 10.1046/j.1365-3113X.1996.10050835.x
- Zhang, X. (1998). Regulatory situation of herbal medicines. A worldwide review. *World Health Organization*, 26, 223-228. Retrieved September 14th 2015 from <http://apps.who.int/medicinedocs/pdf/whozip57e/whozip57e.pdf>