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MEDICINAL IMPORTANCE OF GENUS INULA- A REVIEW

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ABSTRACT

Genus *Inula* is important medicinally as it has anticancerous, antibacterial, hepatoprotective, cytotoxic and anti-inflammatory properties. The species growing in East Asia are widely used by the local people for varied medicinal properties which have prompted many workers to study the phytochemistry of these species which has ultimately resulted in the extraction of a number of novel bioactive molecules. This emphasizes on the need of extensive study for revealing the medicinal importance of the other species of *Inula*.

Keywords: Alkaloids, Bioactive molecules, Himalaya, Medicinal plant, Sesquiterpenes.

INTRODUCTION

The genus Inula belongs to tribe Inuleae, family Asteraceae (Qaiser and Abid, 2003) with about 100 species (Ali et al., 1992). Diverse biological activities have been attributed to this genus, like, anticancer. antibacterial, hepatoprotective, cytotoxic, and anti-inflammatory (Zhao et al., 2006).). As per the available literature the species of reputed medicinal value are distributed in East Asia from Korea to western Pakistan and these include: I. royleana DC. (Kaul, 1997), I. helenium L., I. racemosa Hooker fil., I.viscosa (L.) Aiton, I. britannica L. (Blaschek et al., 1998), I. japonica Thunb.(Jiangsu, 1977), I. rhizocephala var. rhizocephaloids (Clarke) Kitam., I. obtusifolia A. Kern. (Chaurasia et al., 2010), I. grantioids (Kirtikar and Basu, 1933), I. helianthus-aquatica C.Y.Wu (Zeng et al., 2009).

These plants are used in ethno-medicine of the region in which they are growing. These have also been chemically evaluated for their medicinal properties and active principles which reveals the chemical potential of the genus and suggested for further studies in the species which have not been chemically evaluated. The medicinal plants which have been chemically evaluated are as follows:

Inula royleana DC., a medicinal perennial herb, is native to Western Himalaya (Stojakowska and Malarz, 2004). In Kashmir Himalaya it is known as Gugi Phool, is less common and found at an altitude of 2800-3400 m (Khuroo et al., 2007). It is reported that roots of this plant contain sesquiterpene lactones of eudesmane type (Bohlmann et al, 1978; Qurishi et al., 1980), abietane diterpenes (Edwards et al., 1962; Bhat et al., 1975) and diterpene alkaloids (Khaleque et al., 1959; Hegnauer 1964). These compounds possess a variety of biological activities e.g., insecticidal (Jennings et al., 1986), insect repellant (Ulubelen et al., 2001), antimicrobial (Yang et al., 2001), anti-inflammatory (Dirsch et al., 2000) and antiproliferative against different cancer cell lines (Lawrence et al., 2001; Konishi et al., 2002). Some abietanes have vasodepressor effect as well (Kolak et al., 2001; Ulubelen et al., 2002). Roots of this plant are known to have neuromuscular blocking properties (Manchanda et al., 2000) and are used to cure headache (Kala, 2006). In Kashmir Himalaya, shoots of I.

royleana are used for curing dermatitis (Kaul, 1997) and its dried flowers are used for curing throat sores, wounds and inflammation of hooves after boiling in water for half an hour and adding few drops of edible oil (Khuroo et al., 2007). In Pakistan, whole plant is used for intestinal problems (Khan and Khatoon, 2008). Alkaloid of the roots of this plant also helps in lowering hypertension (Haq and Alam, 2010). Roots are stored for protecting garments, its extract is used for curing dermatitis and allergy and root in the form of paste is also applied with leaf on swelling sprains and as an antiseptic in Kangra district of Himachal Pradesh (Prakash and Aggrawal, 2009). Inula helenium L., is a widespread medicinal plant native to Middle Asia, that has been officially listed in some European pharmacopoeias (Blaschek et al., 1998). An essential oil have been reported from the roots of plant containing eudesmane-type this sesquiterpene lactones with alantolactone and isoalantolactone, thymol derivatives (Bohlmann al..1978). triterpenes, sterols et and polysaccharide inulin (Blaschek et al., 1998) forming its main constituents. The sesquiterpene lactones of I. helenium show cytotoxic and antiproliferative activities against human cancer cell lines. Moreover, the compounds also inhibit plant growth (Blaschek et al., 1998; Lawrence et al., 2001; Dirsch et al., 2001; Konishi et al., 2002).

Inula racemosa Hook. f., is a critically endangered Himalayan herb (Anonymous, 1998). It is distributed from temperate to alpine altitudes of Kashmir and Himachal Pradesh where it is known as "Pushkarmoola" and is used as a medicinal plant (Shabir et al., 2010). Its roots are expectorant, seeds are aphrodisiac, roots are used to cure skin diseases and as adultrant for *Sassurea costus* roots (Sarin et al., 1996). Roots of this plant have shown the presence of alantolactones and isoalantolactones (Kalsi et al., 1989; Wang et al., 2000) by chemical profiling. It is due to the presence of these alkaloids that this plant has antifungal properties (Satyawati et al., 1987). This plant is also used by native Americans for treating tuberculosis (Moerman, 1986). Alantolactone present in the roots of *I. racemosa* is useful for reducing fat by enhancing insulin sensitivity (Tripathi and Chaturvedi, 1995) and for treating pulmonary and cardiovascular disorders (Miller, 1998; Lokhande et al., 2006). Its leathery leaves act as sinks for particulate pollutants chiefly carbon particles (Agarwal, 1986).

Inula britannica is a wild plant found in Eastern Asia, including China, Korea, and Japan. The flowers of this plant are used for treating digestive disorders, bronchitis, and inflammation (Bai et al., 2005). Its extracts are reported to have anti-inflammatory, antibacterial, antihepatitic, antidiabetic, and antitumor activities (Jiangsu, 1977; Song et al., 2000; Kobayashi et al., 2002; Song et al., 2002). Various sesquiterpene lactones have been isolated from I. britannica (Zhou et al., 1993; Park and Kim, 1998). Several isolated (Rafi et al., 2005) and derivatized (Liu et al., 2004) sesquiterpene lactones show cytotoxic activity. In addition, kaurane glycosides, as well as some flavonoids and steroids, are reported from I. britannica (Shao et al., 1996).

Inula japonica Thunb., a well known traditional herbal medicinal plant, is widely distributed in Japan, China and Korea (Jiangsu, 1977). The dried leaves and roots of this plant have been used as Chinese folk medicine to treat knife wounds, furunculosis and cough; flowers have peptic, detumescense, antiinflammatory and vermifuge properties (Lin et 1989). Diversified effects such al.. as antidiabetic, hypolipidemia (Han et al., 2004), antifungal, antitumor, antibacterial, hepatoprotective and antihepatitic (Song et al., 2002; Cha et al., 2006 and Wang et al., 2007) have been exhibited to this plant by modern pharmacological study.

Inula viscosa (L.) Aiton (syn. Dittrichia viscosa Greuter), commonly known as "sticky fleabane"

is a perennial weed found in most of the Mediterranean basin (Al-Eisawi, 1998; Baytop, 1999; Al-Dissi et al., 2001). I. viscosa has been used for years in traditional medicine for its antiinflammatory (Barbetti et al., 1985), antipyretic, antiseptic, and antiphlogistic activities (Lauro and Rolih, 1990; Lev and Amar, 2000) and for treating diabetes (Yaniv et al., 1987). It has been found that aqueous extracts of *I. viscosa* possess antifungal activity under in vitro conditions (Qasem et al., 1995; Maoz et al., 1999) with some of its organic solvent extracts showing antibacterial activity (Debat, 1991). Evidence for the antifungal activity in plant extracts was provided by Cohen et al (2002). This herb has also been used in Spanish traditional medicine for treating gastro-duodenal disorders (Lastra et al., 1993). I. viscosa has antiulcerogenic effects (Alkofahi and Atta, 1999), causes abortion (Farnsworth et al., 1975; Karim et al., 1990; Al-Dissi et al., 2001), prevents implantation of zygote in mammals (Al-Dissi et al., 2001), prevents growth of pathogenic fungi (Maoz and Neeman, 2000) and has a strong antioxidant activity (Schinella et al., 2002). There is also evidence for nematicidal/antihelmynth properties of I. viscosa (Oka et al., 2001). I. viscosa contains pharmacologically active compounds some (Ulubelen et al., 1987 and Wollenweber et al., 1991) including sesquiterpenes, sesquiterpene acids (Marongiu et al., 2003), azulenes, lactones, flavonoids, and essential oils (Lauro and Rolih, 1990).

Inula rhizocephala var. *rhizocephaloids* (Clarke) Kitam., an endemic plant species to Nubra and adjacent ranges of Ladakh (Joshi et al., 2006) is a small prostrate herb with yellow flower heads densely clustered at centre and is commonly known as Riamko. It is rare and is found along moist places of North Pullu at an altitude of 4724m. The dried roots of this plant are used for treating colds, cough and chest complaints (Chaurasia et al., 1996). *Inula obtusifolia* A. Kern. is distributed in North-West Himalaya. It is a branched perennial herb with yellow heads, commonly known as Rupmak in Leh valley and found at an altitude of 3353m and is quite useful for domestic animals suffering from diphtheria (Chaurasia et al, 1996). It is also used for tuberculosis, chest problems, cough and as an antiseptic (Khan and Khatoon, 2008).

Inula grantioides, another important medicinal species of *Inula*, has a wide distribution in Sind province. In Pakistan this plant is known as "Naro" and "Kolmur". This plant has a local use in Lasbela for treating patients suffering from asthma (Burdi et al., 1990). It contains an oil that has antibiotic activity (Ahmed and Chughtai, 1961; Kirtikar and Basu, 1933).

Inula helianthus-aquatica C. Y. Wu is a medicinal plant used to treat some cancers in traditional medicine of Yunnan, China. A sesquiterpene lactone namely Bigelovin has been isolated from this herb which acts as an inhibitor to eight cancer cell lines, prominent example being human monoblastic leukemia U937 cells (Zeng et al., 2009).

DISCUSSION

Although there are about 100 species of genus Inula distributed mostly in East Asia but most of the species are growing in the alpine and subalpine areas of the Western Himalaya. Because of the presence of these species in inaccessible areas, only a few species have been subjected to chemical characterisation as evident from perusal of the review of literature during the present study. The species studied clearly indicate that this genus is a potential source for isolation of a variety of chemically defined compounds like sesquiterpene lactones, sesquiterpene acids, triterpenes, sterols, alantolactones, isoalantolactones, flavonoids etc. It is due to the presence of these compounds that the plants of genus Inula act against different diseases like hepatitis, allergy, tumour, ulcers, tuberculosis, cough, colds, diabetes etc. In addition to this,

some species of *Inula* have been exclusively used in ethno-medicine like *Inula grantioides*, *Inula rhizocephala* var. *rhizocephaloids* and *Inula obtusifolia*.

CONCLUSION

of medicinally The synthesis important phytochemicals by the plants of genus Inula has been established beyond doubt. The plant species are being used both in allopathic as well as in traditional systems of medicine for the treatment of a number of ailments. Since, only a few species are being exploited for this purpose, the rest of the species also need to be given due attention as they may prove of immense importance in the production of secondary metabolites and maybe some novel compounds as well.

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REFERENCES

- Agarwal, V.S. (1986) Economic plants of India. Deep printers, 3126, Ramesh Nagar, New Dehli, 187-188.
- Ahmed, M., Chughtai, M.I.D. (1961) Studies on the medicinal plants of Pakistan. Part-I. Pak. J. Sci. & lad. Res. 4: 273.
- 3. Al-Eisawi, D. (1998) Field fuide to wild flowers in Jordan and neighbouring countries. Jordan Foundation Press, Amman, Jordan.
- Al-Dissi, N.M., Salhab, A.S., Al-Hajj, H.A. (2001) Effects of *inula viscosa* leaf extract on abortion and implantation in rats. Journal of Ethnopharmacology **77**: 117–121.

- 5. Ali, S.I., Qaiser, M., Abid, R. (1992) Flora of Pakistan- Asteraceae. University of Karachi, Karachi Printing Press 210: 71.
- 6. Alkofahi. A., Atta, A.H. (1999)Pharmacological screening of the antiulcerogenic effects of some Jordanian medicinal plants in rats. Journal of Ethnopharmacology 67: 341-345.
- 7. Anonymous (1998) Threatened medicinal plants of Himalaya-A check list of CAMP Workshop, Lucknow.
- Bai, N., Lai, C.S., He, K., Zhou, Z., Zhang, L., Quan, Z., Zhu, N., Zheng, Q.Y., Pan, M.H., Ho, C.T. (2005) Sesquiterpene Lactones from *Inula britannica* and Their Cytotoxic and Apoptotic Effects on Human Cancer Cell Lines. J. Nat. Prod. 69: 531-535.
- Barbetti, P., Chiappini, I., Fardella, G., Menghini, A. (1985) A new eudesmane acid from *Dittrichia (Inula) viscosa*. Planta Medica 51: 471.
- Baytop, T. (1999) Therapy with Medicinal Plants in Turkey, Nobel Medical Publication, İstanbul, Turkey.
- Bhat, S.V., Kalyanaraman, P.S., Kohl, H., De-Souza, N.J., Fehlhaber, H.W. (1975) Inuroyleanol and 7-ketoroyleanone, two novel diterpenoids of *Inula royleana* DC. Tetrahedron 31: 1001-1004.
- Blaschek, W., Hansel, R., Keller, K., Reichling, J., Rimpler, H., Schneider, G. (1998) Hagers Handbuchder NY, Folgeband 2:Drogen A-K. pp 523-534.
- Bohlmann, F., Mahanta, P.K., Jakupovic, J., Rastogi, R.C., Natu, A.A. (1978) New sesquiterpene lactones from *Inula* species. Phytochemistry17: 1165-1172.
- Burdi, D.K., Hasan, M., Ahmad, V.U. (1990) Fatty acids of *Inula grantioides*. Pakistan Journal of Pharmaceutical Sciences 3: 33-37.
- Chaurasia, O.P., Singh, B., Sareen, S.K. (1996) Cold Desert Flora Vol. I, Leh valley, Desidoc, Dehli pp 170.
- 16. Cha, M.R., Kim, J.Y., Hwang, J.H. (2006) Cytotoxic activity of the *Inula japonica* extracts

against several human cancer cell lines *in vitro*. Korean J Pharmacogn 37: 130–135.

- Cohen, Y., Baider, A., Ben-Daniel, B.H., Ben-Daniel, Y. (2002) Fungicidal preparations from *Inula viscosa*. Plant Protection Science 38: 629–630.
- 18. Debat, J. (1991) *Inula* extract, its method of preparation and its use in pharmaceutical. US patent no. 4254112.
- Dirsch, V.M., Stuppner, H., Ellmerer Muller, E.P., Vollmar, A.M. (2000) Structural requirements of sesquiterpene lactones to inhibit LPS-induced nitric oxide synthesis in RAW 264.7 macrophages. Bioorg. Med. Chem.8: 2747-2753.
- Edwards, O.E., Feniak, G., Los, M. (1962) Diterpenoid quinines of *Inula royleana* D.C. Can. J. Chem. 40: 1540 – 1546.
- Farnsworth, N.R., Bingel, A.S., Cordell, G.A., Crane, F.A., Fong, H.S. (1975) Potential value of plants as sources of new anti-fertility agents II. Journal of Pharmaceutical Sciences 64: 717– 754.
- 22. Han, M., Wen, J.K., Zheng, B., Zhang, D.Q. (2004) Acetylbritannilactone suppresses NO and PEG2 synthesis in RAW 264.7 macrophages through the inhibition of iNOS and Cox-2 gene expression. Life Sci. 74: 675-684.
- Haq, F., Ahmad, H., Alam, M. (2010) Traditional uses of Medicinal Plants of Nandiar Khuwarr Catchment, District Battagram, Pakistan. Journal of Medicinal Plants Research. 5: 39-48.
- Hegnauer, R. (1964) Chemotaxonomic der Pflanzen. BirkhauserVerlag, Basel. III: 479-480.
- Jennings, K.R., Brown, D.G., Wright, J.R.D.P. (1986) Methyllycaconitine, a naturally occurring insecticide with a high affinity for the insect cholinergic receptor. Experientia 42: 611-613.
- 26. Joshi, P.K., Rawat, G.S., Padilya, H., Roy, P.S. (2006) Biodiversity characterization in Nubra Valley, Ladakh with special reference to plant

resource conservation and bioprospecting. Biodiversity and Conservation 15:4253–4270.

- 27. Jiangsu (1977) Dictionary of Traditional Chinese Materia Medica, Shanghai People's Press: Shanghai 2: 2216-2219.
- Kala, C.P. (2006) Medicinal plants of the high altitude cold desert in India: Diversity, Distribution and Traditional uses. International Journal of Biodiversity Science and Management 2: 43–56.
- Kalsi, S., Goyal, R., Talwar, K.K., Chhabra, B.R. (1989) Stereostructures of two biologically active sesquiterpene lactones from *Inula racemosa*. Phytochemistry 28: 2093-2096.
- Karim, F., Al-Okleh, A., Suleiman, S., Quraan, S. (1990) Poisonous Plants in Jordan, Jordan Natural History Museum, Irbid, Jordan.
- Kaul, M.K. (1997) Medicinal Plants of Kashmir and Ladakh (Temperate and Cold Arid Himalaya). Indus Publishing Company, FS- 5, Tagore Garden, New Dehli pp 126.
- Khaleque, A., Papadopoulos, S., Wright, I., Vento, Z. (1959) Methyl-lycaconitine. Chemistry and Industry (London): 513-514.
- 33. Khan, S.W., Khatoon, S. (2008) Ethnobotanical studies on some useful herbs of Haramosh and Bugrote valleys in Gilgit, Northern areas of Pakistan. Pak.J.Bot. 40: 43-58.
- 34. Khuroo, A.A., Malik, A.H., Dar, A.R., Dar, G.H., Khan, Z.S. (2007) Ethnoveterinary Medicinal uses of some Plant Species by the Gujar Tribe of the Himalaya. Asian Journal of Plant Sciences.6: 148-152.
- 35. Kirtikar, K.R., Basu, B.D. (1933) Indian Medicinal Plants. II: 2nd Edn. pp 1353.
- 36. Kobayashi, T., Song, Q.H., Hong, T., Kitamura, H., Cyong, J.C. (2002) Preventative effects of the flowers of Inula britannica on autoimmune diabetes in C57BL/KsJ mice induced by multiple low doses of streptozotocin. Phytother. Res.16: 377-82.
- Kolak, U., Ari, S., Birman, H., Hasancebi, S., Ulubelen, A. (2001) Cardioactive diterpenoids from the roots of *Salvia amplexicaulis*. Planta Med. 67: 761-763.

- Konishi, T., Shimada, Y., Nagao, T., Okabe, H., Konoshima, T. (2002) Antiproliferative sesquiterpene lactones from the roots of *Inula helenium*. Biol. Pharm. Bull. 25: 1370-1372.
- Lastra, C., Lopez, A., Motilva, V. (1993) Gastroprotection and prostaglandin E2 generation of rats by flavonoids of *Dittrichia viscosa*. Planta Medica 59: 497–501.
- Lauro, L., Rolih, C. (1990) Observations and research on an extract of *Inula viscosa*. Bollettinodella Societa Italiana di Biologia Sperimentale 66: 829-834.
- Lawrence, N.J., McGown, A.T., Nduka, J., Hadf, J.A., Prtichard, R.G. (2001) Cytotoxic Michael-type amine adducts of α-methylene lactones, alantolactone and isoalantolactone. Bioorg. Med. Chem. Lett. 11: 429-431.
- Lev, E., Amar, Z. (2000) Ethnopharmacological survey of traditional drugs sold in Israel at the end of the 20th century. Journal of Ethnopharmacology 72: 191–205.
- 43. Lin, R., Yu, D.J., Wu, Z.Y. (1989) *Inula* L Flora of China. Beijing: Science Press. 263-281
- 44. Liu, S.X., Liu, H., Yan, W., Zhang, L., Bai, N.S., Ho, C.T. (2004). Bioorg.Med. Chem. Lett. 14: 1101-1104.
- Lokhande, P.D., Gawai, K.R., Kodam, K.M., Kuchekar, B.S., Chabukwar, A.R., Jagdale, S.C. (2007) Antibacterial activity of isolated constituents and extract of roots of *Inula racemosa*. Res. J. of Med. Plants. 1: 7-12.
- Manchanda, R., Bhat, S.V., Mehta, B., Karunakaran, J., Venkateswarlu, R. (2000) Alkaloid Extraction of *Inula royleana*. Indian J Physiol Pharmacol. 44: 143-152.
- Maoz, M., Kashman, Y., Neeman, I. (1999) Isolation and identification of a new anti-fungal sesquiterpene lactone from *Inula viscosa*. Planta Medica 65: 281–282.
- Maoz, M., Neeman, I. (2000) Effect of *Inula* viscosa extract on chitin synthesis in dermatophytes and *Candida albicans*. Journal of Ethnopharmacology 71: 479–482.
- 49. Marongiu, B., Piras, A., Pani, F., Porcedda, S., Ballero, M. (2003) Extraction, separation and isolation of essential oils from natural matrices

by supercritical CO_2 . Flavour and Fragrance Journal. 18: 505–509.

- Miller, A.L. (1998) Botanical influences on cardiovascular diseases. Alternative Med. Rev. 3: 422-431.
- Moerman, D.E. (1986) Medicinal plants of native America. University of Michigan AnnArbor., Museum of Anthroplogy. Technical report 19: II.
- 52. Oka, Y., Ben-Daniel, B.H., Cohen, Y. (2001) Nematicidal activity of powder and extracts of *Inula viscosa*. Nematology 3: 735–742.
- Park, E.J., Kim, J. (1998) Cytotoxic sesquiterpene lactones from *Inula Britannica*. Planta Med. 64: 752–754.
- 54. Prakash, V., Aggarwal, A. (2010) Traditional uses of ethnomedicinal plants of lower foothills of Himachal Pradesh-I. Indian Journal ofTraditional Knowledge. 9: 519-521.
- 55. Qaiser, M., Abid, R. (2003) Chemotaxonomic study of *Inula* L.(s.str.) and its allied genera (Inuleae- Compositae) from Pakistan and Kashmir. Pak.J. Bot. 35: 127-140.
- 56. Qasem, J.R., Al-Abed, A.S., Abu-Blan, M.A. (1995) Antifungal activity of clammy *Inula* (*Inula viscosa*) on *Helminthrosporium sativum* and *Fusarium oxysporum* f. sp. *Lycopersici*. Phytopathologia Mediterranea 34: 7–14.
- 57. Qurishi, M.A., Dhar, K.L., Atal, C.K. (1980) A new sesquiterpene lactone from *Inula royleana* roots. Planta Med.38: 282-285.
- Rafi, M.M., Bai, N.S., Ho, C.T., Rosen, R.T., White, E., Perez, D., DiPaola, R.S. (2005) A sesquiterpene lactone from *Inula britannica* induces antitumor effects dependent on Bcl-2 phosphorylation. Anticancer Res. 25: 313-318.
- Sarin, Y.K. (1996) Illustrated Manual of Herbal Drugs used in Ayurveda. National Institute of Science Communication (CSIR), Dr. K. S. Krishnan Marg. New Dehli, India.
- Satyawati, G.V., Gupta, A.K., Neeraj, T. (1987) Medicinal Plants of India. Indian Council of Medical Research, New Dehli.
- 61. Schinella, G.R., Tournier, H.A., Prieto, J.M., Mordujovich, P., Rios, J.L. (2002) Antioxidant
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activity of anti-inflammatory plant extracts. Life Sciences 70: 1023–1033.

- 62. Shabir, P.A., Nawchoo, I.A., Wani, A.A. (2010) Development of vegetative and sexual multiplication protocol for commercialization of *Inula racemosa* Hook.f., a critically endangered medicinal plant of N.W. Himalaya. Nature and Science. 8: 246.
- 63. Shao, Y., Bai, N.S., Zhou, B.N. (1996) Kaurane glycosides from *Inula britannica*. Phytochemistry 42: 783-786.
- 64. Song, Q.H., Kobayashi, T., Iijima, K., Hong, T., Cyong, J.C. (2000) Hepatoprotective effects of *Inula britannica* on hepatic injury in mice. Phytother. Res. 14: 180-186.
- Song, Q.H., Kobayashi, T., Hong, T., Cyong, J.C. (2002) Effects of *Inula britannica* on the production of antibodies and cytokines and on T cell differentiation in C57BL/6 mice immunized by ovalbumin. Am. J. Chin. Med. 30: 297-305.
- 66. Stojakowska, A., Malarz, J. (2004) *In vitro* propagation of *Inula royleana*. Acta Societatis Botanicorum Poloniae 73: 5–8.
- Tripathi, Y.B., Chatturvedi, P. (1995) Assessment of endocrine response of *Inula racemosa* in relation to glucose homeostasis in rats. Ind. J. Exp. Biol. 33: 686-689.
- Ulubelen, A., Birman, H., Oksuz, S., Topcu, G., Kolak, U., Barla, A., Voelter, W. (2002) Cardioactive diterpenes from the roots of *Salvia eriophora*. Planta Med. 68: 818-821.
- Ulubelen, A., Mericli, A.H., Mericli, F., Kilincer, N., Ferizli, A.G., Emekci, M., Pelletier, S.W. (2001) Insect repellent activity of diterpenoid alkaloids. Phytother. Res. 15: 170-171.
- Ulubelen, A., Öksüz, S., Gören, N. (1987) Sesquiterpene acids from *Inula viscosa*. Phytochemistry 26: 1223–1224.

- Wang, C.M., Jia, Z.J., Zheng, R.L. (2007) The effects of 17 sesquiterpenes on cell viability and telomerase activity in the human ovarian cancer cell line HO- 8910 [J]. Planta Med. 73: 180-184.
- 72. Wang, K., Liu, H., Zhao, Y., Chen, X., Hu, Z., Song, Y., Ma, X. (2000) Separation and Determination of alantolactone and isoalantolactone in traditional Chinese herbs by capillary electrophoresis. Talanta.52: 1001-1005.
- 73. Wollenweber, E., Mayer, K., Roitman, J.N. (1991) Exudate flavonoids of *Inula viscosa*. Phytochemistry. 30: 2445–2446.
- 74. Yang, Z., Kitano, Y., Chiba, K., Shibata, N., Kurokawa, H., Doi, Y., Arakawa, Y., Tada, M. (2001) Synthesis of variously oxidized abietane diterpenes and their antibacterial activities against MRSA and VRE. Bioorg. Med. Chem. 9: 347-356.
- Yaniv, Z., Dafni, A., Friedman, L., Palevitch, D. (1987) Plants used for the treatment of diabetes in Israel. Journal of Ethnopharmacology. 19: 145–151.
- 76. Zeng, G.Z., Tan, N.H., Ji, C.J., Fan, J.T., Huang, H.Q., Han, H.J., Zhou, G.B. (2009) Apoptosis Inducement of Bigelovin from *Inula helianthus-aquatica* on Human Leukemia U937 Cells. Phytother. Res. 23: 885–891.
- Zhao, Y.M., Zhang, M.L., Shi, Q.W., Kiyota, H. (2006) Chemical constituents of plants from the genus *Inula*. J. Chem. Biod. 3: 371–384.
- Zhou, B.N., Bai, N.S., Lin, L.Z., Cordell, G.A. (1993) Sesquiterpene lactones from *Inula britannica*. Phytochemistry 34: 249-252.