

Rec. Nat. Prod. 4:1 (2010) 1-25

records of natural products

Chemical Constituents and Biological Activities of Artemisia herba-alba

Abou El-Hamd H. Mohamed^{1*}, Magdi. A. El-Sayed², Mohamed E. Hegazy³,

Soleiman E. Helaly¹, Abeer M. Esmail¹ and Naglaa S. Mohamed¹

¹ Department of Chemistry, Aswan-Faculty of Science, South Valley University, Aswan, Egypt ²Department of Botany, Aswan-Faculty of Science, South Valley University, Aswan, Egypt ³Chemistry of Medicinal Plant Department, National Research Centre, Dokki, Giza, 12622,

Egypt

(Received June 7, 2009; Revised November 18, 2009; Accepted November 22, 2009)

Abstract: Artemisia, one of the larger genera in the family Asteraceae and the largest genus in the tribe Anthemideae, comprises from 200 to more than 500 taxa at the specific or subspecific level. Many Artemisia species have a high economic value in several fields, as food plants and as antihelminthic and antimalaria in medicine. Artemisia herba-alba was known for its therapeutic and medicinal properties, it was used in both traditional and modern medicine. Several papers have been published on the chemical composition of specimens of A. herba-alba. The aim of this work is to review all available scientific literature published on A. herba-alba. The focus will be on the chemical constitutions which have been identified from this species, in addition to all of the reported biological activities of this species have been included as well as the pharmacology and toxicology.

Keywords- Artemisia herba-alba; sesquiterpenes; flavonoids; essential oil; biological activities.

1. Introduction

The genus Artemisia L. (family Asteraceae, tribe Anthemideae), comprises a variable number of species (from 200 to over 400, depending on the authors) found throughout the northern half of the world. The genus may be divided into sections Artemisia and Dracunculus [1]. The genus Artemisia is known to contain many bioactive compounds; artemisinin exerts not only antimalarial activity but also profound cytotoxicity against tumor cells [2] and arglabin is employed for treating certain types of cancer in the former USSR [3]. Over the past decade Artemisia species have been used traditionally in varies populations, thus; A. keiskeana Miq has been used as a Traditional Chinese drug for the treatment of gynaecepathy, amenerrhea, bruise and rheumatic disease [4], A. vestita Wall ex DC. has been utilized for the treatment of fungal infections such as tinea, tympanitis, and thrush [5], A. abrotanum L. was found to possess spasmolytic activity on the

^{*} Corresponding author: E-Mail: abuelhamd2002@yahoo.com, Phone: +2-972312850. Fax: +2-973480450

carbacholine induced contraction of guinea-pig trachea [6]. *Artemisia* species are popular plants which are used for the treatment of diseases such as hepatitis, cancer, inflammation and infections by fungi, bacteria, and viruses [7]. Furthermore, several species of *Artemisia* are used in folk medicine, thus; *A. vulgaris* has been employed in the treatment of painful menstruation and in the induction of labor or miscarriage [8], *A. mongolica* Fisch has been used as a folk medicine for generations to cure inflammations and colds in northwest China [9], *A. pontica* L. is well known in Bulgarian folk medicine as a sedative and an appetizer [10].

Artemisia herba-alba Asso, known also as desert wormwood (known in Arabic as shih, Armoise blanche (Fr.)) [11], has been used in folk medicine by many cultures since ancient times, used in Moroccan folk medicine to treat arterial hypertension and/or diabetes [12-14]. Herbal tea from this species has been used as analgesic, antibacterial, antispasmodic, and hemostatic agents [15]. During an ethnopharmacological survey carried out among the Bedouins of the Negev desert, it was found that *Artemisia herba-alba* relieved stomach disorders [16]. This plant is also suggested to be important as a fodder for sheep and for livestock in the plateau regions of Algeria where it grows abundantly [17-19]. Ascaridae from hogs and ground worms were killed by the oil of the Libyan *A. herba-alba* in a short time [20, 21].

2. Botanical aspects

2.1 Morphology

A. herba-alba is a greenish-silver perennial herb grows 20-40 cm in height; it is a chamaeophyte (i.e. the buds giving rise to new growth each year are borne close to the ground). The stems are rigid and erect. The grey leaves of sterile shoots are petiolate, ovate to orbicular in outline whereas leaves of flowering stems are much smaller. The flowering heads are sessile, oblong and tapering at base. The plants flower from September to December. Plants are oblong and tapering at base. Plants are found on the steppes of the Middle East and North Africa where they are common and sometimes stand-forming (Figure 1) [22].



Figure 1. Photograph of Artemisia herba-alba.

2.2 Taxonomy

Artemisia herba-alba is classified into Kingdom: Plantae, Subkingdom: Tracheobionta, Superdivision: Spermatophyta, Division: *Magnoliophyta*, Class: Magnoliopsida, Subclass: Asteridae, Order: Asterales, Family: Asteraceae, Subfamily: Asteroideae, Tribe: Anthemideae, Subtribe: Artemisiinae, Genus: Artemisia L., subgenus: Seriphidium and species: *Artemisia herba-alba* Asso [23-25].

2.3 Distribution

The genus *A. herba-alba* is a medicinal and aromatic dwarf shrub that grows wild in arid areas of the Mediterranean basin, extending into northwestern Himalayas. This plant is abundant in the Iberian Peninsula and reaches highest population in the centre of Spain spreading over the eastern, southeastern and southern Spain. This taxon grows wild on nitrofilous and gypsum-rich substrata (Figure 2) [26, 27].



Figure 2. Distribution map of Artemisia herba-alba

3. Phytochemistry

Various secondary metabolites have been isolated from *A. herba-alba*, perhaps the most important being the sesquiterpene lactones that occur with great structural diversity within the genus *Artemisia*. Additional studies have focused on flavonoids and essential oils [28].

3.1 Sesquiterpene lactones

Sesquiterpene lactones are among the prominent natural products found in *Artemisia* species and are largely responsible for the importance of these plants in medicine and pharmacy. Several structural types of sesquiterpene lactones were found in the aerial parts of *A. herba-alba*. Eudesmanolides followed by germacranolides seem to be the most abundant types of lactones found in this species.

In course of investigation of *A. herba-alba* plants growing in Israel (Negev and Judean desert) five different chemotypes were identified on the basis of differences in their sesquiterpene lactone constitution (1-10) (Figure 3) [29-34]. In Spain, some phytochemical studies have investigated the sesquiterpene lactones from the *A. herba-alba*, (11-47) collected in different geographical location. (Figure 3) [35-41].

Many groups have studied the chemistry of *A. herba-alba* growing in Egypt. Most studies have been concerned with the sesquiterpene lactones. All the lactones isolated (**48-57**) differ from those found previously from *A. herba-alba* growing in Israel (Fig. 3) [42, 43]. Few literature papers studied the chemical constituent of Moroccan *A. herba-alba* species (**58-64**) [44, 45], and the Algerian one (**65-66**) [15], which proven that this genus is rich in sesquiterpenes (Figure 3).

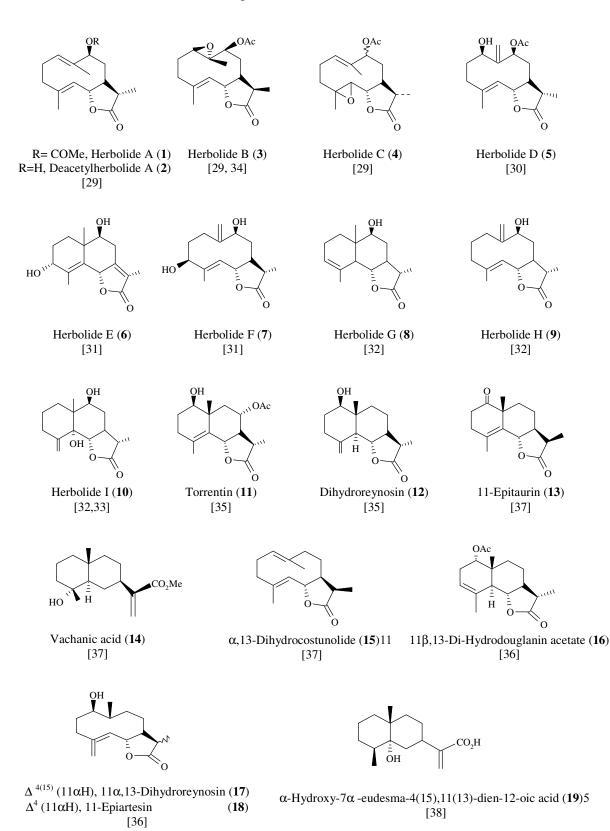
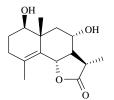
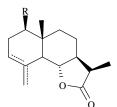
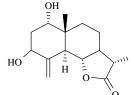


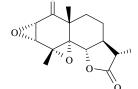
Figure 3. Sesquiterpene lactones from A. herba-alba growing in Egypt, Morocco, Spain, Algeria and Israel



1 β ,8 α -Dihydroxyeudesm-4-en-6 β ,7 α ,11 β H-12,6-olide (**20**) [36]

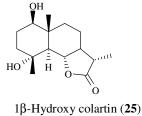






 $1-Oxo-2\alpha, 3\alpha, 4\alpha, 5\alpha$ -diepoxyeudesman- 11β H-12, 6\alpha-olide (24)

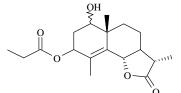
[39]



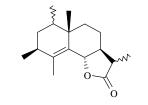
[38]

3-Epi-erivanin (**23**) [38]

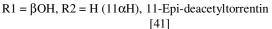


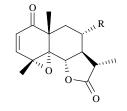


1β-OH, 1β-Hydroxy-3β-propionyloxy- $6\beta,7\alpha,11\beta$ H-eudesm-4-en-12,6-olide (29) 1α-OH, 1α-Hydroxy-3β-propionyloxy- $6\beta,7\alpha,11\beta$ H-eudesm-4-en-12,6-olide (30) [38]

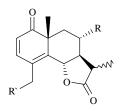


 $\begin{array}{ll} \text{R1} = \beta \text{-OH}, \text{R2} = \text{Ac} (11\alpha\text{H}), 11\text{-Epitorrentin} & \textbf{(31)} \\ \text{R1} = \alpha \text{-OH}, \text{R2} = \text{Ac} (11\alpha\text{H}), 1, 11\text{-Diepitorrentin} & \textbf{(32)} \\ \text{R1} = \beta \text{-OH}, \text{R2} = \text{H} (11\beta\text{H}), \text{Deacetyltorrentin} & \textbf{(33)} \\ \text{R1} = \beta \text{OH}, \text{R2} = \text{H} (11\alpha\text{H}), 11\text{-Epi-deacetyltorrentin} & \textbf{(34)} \end{array}$



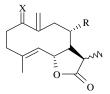


 $R = H, 1-Oxo-4\alpha, 5\alpha-Epoxyeudesm-2$ -en-11 β H-12,6 α -olide (35) $R = OH, 1-Oxo-8\alpha-Hydroxy-4\alpha, 5\alpha$ epoxyeudesm-2-en-11 β H-12,6 α -olide (36) [39]

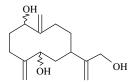


$R = R^{*} = H (11\alpha H)$, 1-Oxoeudesma-2,4-dien	
-11\alphaH-12,6\alpha-olide	(37)
$R = R^{\sim} = H (11\beta H), 1$ -Oxoeudesma-2,4-dien	
-11βH-12,6α-olide	(38)
$R = OH, R^{\sim} = H (11\alpha H), 1-Oxo-8\alpha$ -hydroxy-	
eudesma-2,4-dien-11\alphaH-12,6\alpha-olide	(39)
$R = OH, R^{T} = H (11\beta H), 1-Oxo-8\alpha-hydroxy-$	
eudesma-2,4-dien-11βH-12,6α-olide	(40)
$R = H, R^{\sim} = OH (11\beta H), 1-Oxo-15-hydroxy-$	
eudesma-2,4-dien-11βH-12,6α-olide	(41)

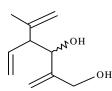




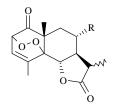
$$\begin{split} R &= H, X = O (11\alpha H), 1\text{-}Oxogermacra-4,10(14)-\\ & \text{dien-}6\beta,7\alpha,11\alpha H\text{-}12,6\text{-}olide (45)\\ R &= OH, X = H, \beta OH (11\alpha H), 11\text{-}Epishonachalin A (46)\\ R &= OH, X = H, \beta OOH (11\beta H), 1\beta\text{-}Hydroperoxy-8\alpha\\ & \text{-}hydroxygermacra-4,10(14)\text{-}dien-6\beta,7\alpha,11\beta H\text{-}12,6\text{-}olide (47)\\ & [36] \end{split}$$



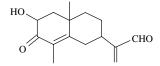
1β,5,12-Trihydroxygermacra-1(10), 4(15), 11(13)-triene (**49**) [43]



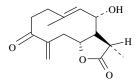
7-Hydroxy-5,6-dehydro-4,5-dihydrolyratrol (**51**) [43]



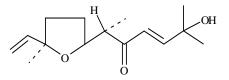
$$\begin{split} R &= H, (11\alpha H), 1\text{-}Oxo\text{-}2\alpha,5\alpha\text{-}peroxyeudesm} \\ &-3\text{-}en\text{-}11\alpha H\text{-}12,6\alpha\text{-}olide} \\ R &= H (11\beta H), 1\text{-}Oxo\text{-}2\alpha,5\alpha\text{-}peroxyeudesm} \\ &-3\text{-}en\text{-}11\beta H\text{-}12,6\alpha\text{-}olide} \\ R &= OH (11\beta H), 1\text{-}Oxo\text{-}8\alpha\text{-}Hydroxy\text{-}2\alpha,5\alpha} \\ &-peroxyeudesm\text{-}3\text{-}en\text{-}11\beta H\text{-}12,6\alpha\text{-}olide} \\ \end{split}$$



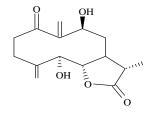
 2β -Hydroxy-13-oxo- α -cyperene **48**) [43]



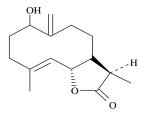
 3β -Hydroxy-8-oxo-6 β H,7 α H,11 β Hgermacran-4(14),9(10)-dien-6,12-olide (**50**) [42]

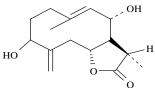


2,6,10-Trimethyl-Cis-7,10-oxido-dodeca-3E,11-dien-2-ol-5-one (**52**) [42]



 5β ,9 β -Dihydroxy-1-Oxo-germacra-1(10),4(15)-dien-12,6-olide (55) [43]





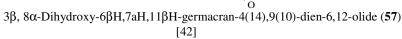
1β-Hydroxy-6βH,7αH,11αH-germacran-4(5)-10(15)-dien-6,12-olide (**56**) [42]

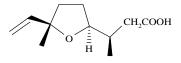
0



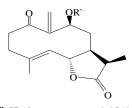
oш

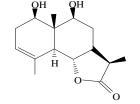
0





(3 R,4S,7R)-3,7-Dimethyl-4,7-epoxynon-8-enoic acid (**59**) [44]

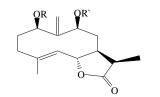




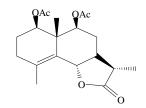
Herbalbin (**58**) [45]

1 β ,9 β -Dihydroxyeudesm-3-en-5 α ,6 β ,11 β H-12,6-olide (60) [44]

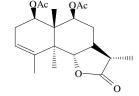
Figure 3. Continued



 $R = R^{*} = H, Deacetylherbolide D (63)$ $R = H, R^{*} = Ac, 1b-Hydroxy-9b-acetoxygermacra
-4,10(14)-dien-6b,11bH-12,6-olide (64)
[44]$



1β,9β-Diacetoxyeudesm-4-en-6β,11bH-12,6-olide (**65**) [15]



1β,9β-Diacetoxyeudesm-3-en-6β,11βH-12,6-olide (**66**) [15]

Figure 3. Continued

3.2 Flavonoids

The flavonoids detected in *A. herba-alba* show a large structural variation, ranging from common flavone and flavonol glycosides to more unusual highly methylated flavonoids. In studies of the leaves and steams of *A. herba-alba* collected from Sinai, a total of eight flavonoids *O*- and *C*-glycoside were isolated and identified [46, 47]. Examination of the aerial parts of *A. herba-alba* collected from Lebanese herbal stores led to the isolation of two flavonoids; hispidulin and cirsilineol [48] (Fig. 4). A new flavone, 5,4'-dihydroxy-6,7,3'-trimethoxyflavone, was isolated from the nonglycosidic extract of the aerial parts of *A. herba-alba* [49].

4. Phenolic compounds & waxes

Chlorogenic acid was observed in *A. herba-alba*, when a chemical survey of 49 Moroccan medicinal plant species was performed by ESR spectroscopy [50]. During a survey for antiulcerogenic principles of *A. herba-alba*, eight polyphenolics and related constituents were isolated. These included chlorogenic acid, 4,5-*O*-dicaffeoylquinic acid, isofraxidin 7-*O*- β -D-glucopyranoside, 4-*O*- β -D-glucopyranosylcaffeic acid, rutin, schaftoside, isoschaftoside, and vicenin-2 [51]. In a study of the components of *A. herba-alba* wax, obtained in 0.23% yield by extraction of the dry plant with ether, contained 32.1% saturated C16-32 acids (35.2% C28 and 26.5% C30), 23.2% saturated. C21-31 hydrocarbons (67.7% C29 and 24.2% C31), 27.1% esters (mainly of saturated C18, C19, and C20 acids and saturated C22 and C24 alcohols.), and 16.96% saturated C16-26 alcohols. (C16 24.71%, C20 10.34%, C22 32.88%, and C24 22.96%) [52].

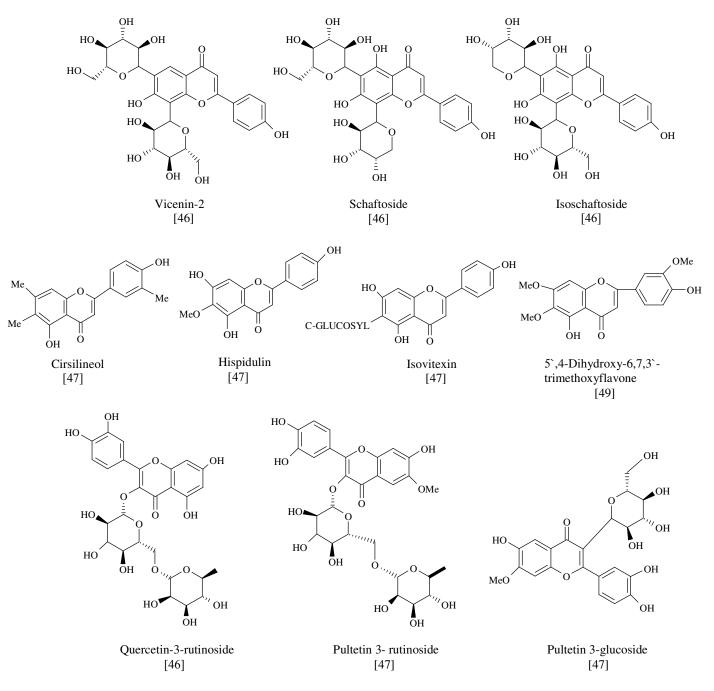


Figure 4. Flavonoids Artemisia herba-alba.

5. Essential oils

Over the last decades, the *Artemisia herba-alba* oil; called scheih oil [53], has been thoroughly investigated and the diversity in oil composition from plants grown in different countries and even those from different localities in the same country have led to the many oil-dependent chemotypes assigned to the plant [54]. Generally, the oil was largely reported to be composed of monoterpenoids, mainly oxygenated, such as 1,8-cineole, chrysanthenone, chrysanthenol (and its acetate), α/β -thujones, and camphor as the major components [55, 56].

High variability of volatile constituents was observed when different populations of *A. herba-alba* collected at various sites in Israel were compared [57]. Whereas samples of *A. herba-alba* collected at Elat contained chrysanthenyl acetate as major component (31%) followed by chrysanthenol (6.4%) and the acetophenone xanthocyclin, the essential oil of *A. herba-alba* from the Judean desert exhibited 1,8-cineole as the major compound (50%) followed by appreciable amounts of α - and β -thujone (27%) and other oxygenated monoterpenes such as terpinen-4-ol (3.3%), camphor (3%) and borneol (3%) [58, 59]. Essential oils of *A. herba-alba* collected at other localities in Israel contained some more unusual volatile terpenes including artemisia alcohol and lyratol [60]. Also, Two oil types were found for plants grown in Israel and Sinai, those of the cineol-thujane-bornane type and the pinane type; the oil of all studied populations contained 1,8-cineole in varying concentrations. More recently, it was reported that a further five chemotypes could be distinguished in plants growing in the Holy Land and Sinai due to the variation on the major components of the oil, suggesting the existence, in this region, of a greater number of chemovarieties than previously believed [61]. In addition, artemisia ketone was reported as the principle component of an Egyptian chemotype [62], while a French type [63] was found to predominate in 1,8-cineole, camphor and chrysanthenone.

In Jordan, regular monoterpenes were predominant (39.3%) and the principal components were α - and β -thujones (27.7%). The other major identified components were: sabinyl acetate (5.4%), germacrene D (4.6%), α -eudesmol (4.2%) and caryophyllene acetate (5.7%) [64].

In Morocco, the oil there was generally characterized by substantial levels of ketones such as α - and β thujones and camphor [65, 66, 67], whereas davanone and/or chrysanthenyl acetate were the major oil components in other chemotypes [68]. In the market leader of *A. herba-alba* essential oil exports, 16 chemotypes were found [69], with 12 having monoterpenes as major components and for four sesquiterpene skeletons represent the major fraction of the oil, investigations reported no correlations between chemotypes and geographic distribution [70].

Studies from Spain showed that monoterpene hydrocarbons and oxygenated monoterpene are the most abundant skeletons in *A. herba-alba* oil, but large amounts of sesquiterpenes were found for some populations. Camphor, 1,8-cineole, *p*-cymene and davanone were the major components found [71, 26]. Another study from Spain showed that the chemotaxonomic affinity between Spanish and Israeli populations of the same plant was not reflected in their oil compositions. The Spanish oil contained large amounts of sesquiterpenes but lacked significant quantities of thujane derivatives. However, a more recent investigation from Spain showed the sesquiterpene davanone to be the principal component of the oil, which was also dominated by the *p*-menthane and pinane skeletons [72, 73, 74,75].

In Tunisian oil oxygenated monoterpenes were found to be the major components of *A. herba-alba* oil extracted from aerial parts of plants originated from arid regions [76, 77]. In another study of the Tunisian *Artemisia herba-alba* oil, the main components were cineole, thujones, chrysanthenone, camphor, borneol, chrysanthenyl acetate, sabinyl acetate, davana ethers and davanone. Monoterpenes, sesquiterpenes are found in some samples as major components. The chemical compositions revealed that some samples have compositions similar to those of other *Artemisia herba-alba* essential oils analyzed in other countries [78].

In an Algerian oil, camphor, α/β -thujones, 1,8-cineole and chrysanthenyl derivatives were the major components [79, 80, 81]. In another studies, camphene (3%), borneol (3.6%), davana ether (8.8%), davanone (36.1%) were the major components [82-88]. The essential oil obtained from the aerial parts of *A. herba-a*lba growing wild in M'sila-Algeria, contained camphor (19.4%), trans-pinocarveol (16.9%), chrysanthenone (15.8%) as major components. Monoterpenoids are the main components (86.1%) and the

irregular monoterpenes (3.1%) [89]. On the other hand, other components have previously been found in other *A. herba-alba* oils such as (*Z*)-jasmone, xanthoxylin were not detected in this oil [90, 91].

One study of *A. herba-alba* in Cyrenaica (Libya) had reported that the dried grass contained 0.29% of an essential oil containing 6.7% cineole [92].

6. Biological effect of A. herba-alba

6.1 Antioxidant activity

Many medicinal plants contain large amounts of antioxidant compounds, which could be isolated and then used as antioxidants for the prevention and treatment of free radical-related disorders. In a study by Djeridane [93], the purpose was the evaluation by a chemical method of the antioxidant capacity of phenolic compounds in some Algerian medicinal plants, including *A. herba-alba*. These medicinal plants showed stronger antioxidant activity and content in phenolics than the common nutritional plants. It has been also noted in this study that these Algerian plants are strong radical scavengers and can be considered as good sources of natural antioxidants for medicinal and commercial uses [93]. Raw and cooked ground beef patties were treated with an aqueous extract of *A. herba-alba*, rosemary, fennel and rue at levels of 5 mm of 10% (w/w vegetable matter to water) extract for every 100 g of meat. Patties were kept under refrigeration (4°) for a period of 16 days, and samples were drawn at 4-day intervals. Results showed that cooked meat was more susceptible to oxidative deterioration than raw meat. In addition, *A. herba-alba* had a somewhat less effective role than the other herbs [94]. In another study, 21 plant samples were collected from different Jordanian locations and used for antioxidant evaluation. The level of antioxidant activity, determined by DPPH and ABTS assays, showed that *Artemisia herba-alba* has a moderate antioxidant activity compared to the other plants [95].

Abid compare the long-term effects of Artemisia herba-alba decoction with a green or black tea decoction, prepared without sugar, on the antioxidant processes in rats. The conclusion of this study showed that Artemisia, as well as green tea decoctions, increased the total antioxidant status, whole blood glutathione peroxidase activity and zinc and copper status, and prevented weight gains and increased conjugated dienes, plasma glucose, lipids and iron status. The beneficial antioxidant effects were in descending order: Artemisia decoction \geq green tea decoction > black tea decoction. So, Artemisia could constitute a good adjuvant to combat obesity, hyperglycemia, hyper-triglyceridemia, hyper-cholesterolemia and particularly oxidative stress [96].

The effects of seven medicinal plants including *Artemisia herba-alba* on protein degradation, lipid peroxidation, erythrocyte deformability and osmotic fragility of erythrocytes exposed in vitro to 10 mM H_2O_2 for 60 min at 37°C have been examined. The result was that *Artemisia herba-alba* did not protect erythrocytes against lipid peroxidation [97].

6.2 Anti-venom activity

Aqueous extracts of 12 medicinal plants traditionally used in Jordan for the inhibition in humans of snake and scorpion venoms were evaluated for their possible anti-venom activity. Among the plants tested, 9 extracts were found to inhibit the hemolytic activities of both venoms. The most active plant extract was *Artemisia herba-alba*, which gave 100% inhibition [98].

6.3 Antifungal activity

The antifungal activity of *Artemisia herba-alba* was found to be associated with two major volatile compounds isolated from the fresh leaves of the plant. Carvone and piperitone were isolated and identified by GC/MS, GC/IR, and NMR spectroscopy. Antifungal activity was measured against *Penicillium citrinum* (ATCC 10499) and *Mucora rouxii* (ATCC 24905). The antifungal activity (IC_{50}) of the purified compounds carvone and piperitone was estimated to be 5 µg/ml and 2 µg/ml against *Penicillium citrinum*, and 7 µg/ml and 1.5 µg/ml against *Mucora rouxii*, respectively [99]. In another study, the antifungal activity of the

12

essential oils of 25 Moroccan medicinal plants, including A. herba-alba, against Penicillium digitatum, Phytophthora citrophthora, Geotrichum citri-aurantii, and Potrytis cinerea. A. herba-alba essential oil showed only weak antifungal activity at 250 µg/ml concentration [100]. In addition, the effect of A. herbaalba, Eucalyptus, and Rosmarinus essential oils was evaluated on the mycelial growth and toxigenesis of Penicillium aurantiogriseum and P. vindication. A significant decrease in mycelial dry weight was obtained with the addition of 0.05-2.5% of each of the three essential oils in yeast extract sucrose broth. The inhibition of mycelium growth was tested on malt extract agar, Czapeck yeast agar, yeast extract sucrose agar and broth at constant pH, and was highly effective for A. herba-alba, followed by Eucalyptus. A complete inhibition of toxin production was observed with 0.44% of each essential oil for P. aurantiogriseum and 0.22% for P. viridicatum [101]. The effect of A. herba-alba and Oreganum oils on spore germination, mycelial elongation and sporulation were studied in three fungi. All three stages of fungal asexual reproduction were affected but mycelium growth was the most sensitive, followed by spore germination and then sporulation of the three fungi studied. Zygorrhynchus sp. was found to be the most sensitive followed by Aspergillus niger and then Penicillium italicum. A. herba-alba was less active on the three phenomena studied than the Oreganum oil. [102]. In addition, A. herba-alba essential oil demonstrated a synergistic action on the inhibition of mycelium growth in Zygorrhynchus sp. and Aspergillus niger isolates, when associated with sodium chloride or fatty acid [103]. The essential oils extracted from 10 Algerian plants; including A. herba-alba, were analyzed for their potential activity against Candida albicans. A moderate efficiency was obtained with the essential oil from A. herba alba which showed an antifungal effect 5617-fold lower than that measured with amphotericin B. [104].

The inhibitory effects of *Artemisia* flower heads (*Artemisia herba-alba*), on the growth and aflatoxin production of a toxigenic strain of *Aspergillus flavus* was tested using different concentrations. The plant inhibited aflatoxin formation by 85-90% of that of control at a concentration of 10% [105].

The *in-vitro* antifungal activity of *A. herba-alba* essential oil has been evaluated on various microorganisms. The oil showed a very strong action vs. *Candida* and *Microsporum*. [106]

6.4 The nematicidal activity

The *in-vitro* nematicidal activity of methanolic extracts (20 µg/ml) from twenty Jordanian plant species against two species of root-knot nematodes was evaluated. The leaf extract of *Artemisia herba-alba* was the most effective causing 22, 51, and 54% mortality after 24, 48 and 72 h of exposure, respectively [107].

6.5 Antibacterial activity

The antibacterial activity of Artemisia herba-alba collected near Sde-Boker (Negev desert), Israel, has been investigated. Only the essential oil was found to be active against some Gram-positive bacteria (Streptococcus hemolyticus and Staphylococcus aureus) and Gram-negative bacteria (Escherichia coli, Shigella sonnei and Salmonella typhosa,). The essential oil was fractionated by column chromatography, and these fractions were tested for antibacterial activity. The principal component of the most active fraction was santolina alcohol [108]. Furthermore, the essential oils from four Artemisia herba-alba populations collected in Israel (Sde-Boker, Mizpe Ramon, Judean desert and Elat) were investigated for their antibacterial activity. All the oils had slight antibacterial activities in the concentration range of 1-2 mg/ml. The oils were active against gram negative bacteria (Escherichia coli, Shigella sonnei, Salmonella typhosa, Serratia marcescens and Pseudomonas aerugenosa) and against gram positive bacteria (Bacillus subtilis, Streptococcus hemolyticus and Staphylococcus aureus). The oil derived from the Sde Boker variety exhibited the highest antibacterial activity, especially against the Streptococcus, Pseudomonas and Serratia strains tested. This oil, as well as those from Mizpe Ramon and the Judean Desert, showed relative high activity against S. sonnei and S. typhosa whereas they were almost inactive against E. coli. The oil derived from plants collected near Elat possessed very low activities. The antibacterial effect may explain the extensive use of A. herba-alba in folk medicine [57,109]. Also, the in vitro antibacterial activity of A. herba-alba essential oil has been evaluated on virous microorganisms, The oil showed a very strong action vs. *Staphylococcus*, The inhibiting action of the oils was low vs. the *enterobacteria* [106]. The aqueous extract of *A. herba-alba* possessed relatively weak antibacterial activity and virtually little or no inhibitory activity against the yeast *Saccharomyces cerevisiae* [110]. The antibacterial activity of *A. herba-alba* performed against *Bacillus subtilis* and *Escherichia coli* showed no significant activity against either species [111].

Mycoplasmas are one of the smallest free-living microorganisms. Unlike other bacteria they lack a rigid cell wall. Hence, they are not susceptible to penicillins and other antimicrobials that act on this structure. *Artemisia herba-alba*, was one among six Jordanian traditional medicinal plant methanolic extracts that were tested against 32 isolates of *Mycoplasma* species. The result of this study showed that the most effective plant extract in vitro against all *Mycoplasma* species was *A. herba-alba* with MIC values of 3.125-6.25 mg/ml. Therefore, this plant could be considered as a readily available alternative for drugs such as the fluoroquinolones, tetracyclines, macrolides and chloramphenicols that are currently used in the treatment of *Mycoplasma* infections [112].

6.6 Antispasmodic activity

The essential oils from four *Artemisia herba-alba* populations collected in Israel (Sde-Boker, Mizpe Ramon, Judean desert and Elat) were investigated for their antispasmodic activity. The *A. herba alba* oils were tested on isolated rabbit jejunum. The results indicate that the different oils showed similar relaxation patterns. The concentrations, which induced 50% relaxation were similar for the Sde Boker, Mizpe Ramon and Judean Desert oils (ca 3x10-5%), while that of the Elat oil was ten times lower [57].

6.7 Anthelmintic activity

The anthelmintic activity of the powdered shoots of *Artemisia herba-alba* was investigated in experimental haemonchosis in six Nubian goats which had been infected with single doses of 800 to 1,000 infective *Haemonchus* larvae. The clinical signs of caprine haemonchosis included inappetence, dullness and soft faeces and were correlated with the pathological findings. None of these changes were observed in four of the six goats following the treatment with 2, 10 or 30 g of *Artemisia* shoots. This successful therapy was supported by the absence of eggs in the faeces or adult worms in the abomasum at necropsy and of significant lesions in the tissues of the goats and return of the concentrations of serum ammonia, sodium, potassium, total protein and creatinine and of the activity of aspartate aminotransferase (GOT) to normal. In two goats, treated with either 10 or 30 g of *Artemisia* shoots, egg production was not completely suppressed and a few adult *Haemonchus* worms were found in the abomasum [113, 109].

Al-Waili suggested that the aqueous extract of *A. herba-alba* might have a possible therapeutic value in intestinal infection with *Enterobius vermicularis*. Therefore the possible effect of *A. herba-alba* extract on *E. vermicularis* infection was examined in 10 patients. The results appeared to show that *A. herba-alba* extract eradicated intestinal infection with *E. vermicularis* within 3 days in all 10 patients treated [114].

6.8 In vitro Antileishmanial activity

The aqueous extract and essential oil of *Artemisia herba-alba* were tested for their antileishmanial activity against *Leishmania major*. The strongest leishmanicidal activity was observed with the essential oil at 2 μ g/ml compare to the other two strains tested. The aqueous extract showed an antileshmanial activity at concentration of 4 μ g/ml [115].

6.9 Neurological activities

In the recent decades the use of traditional medicine in Lebanon has increased. Aqueous, ethanol and ethyl acetate extracts of seven Lebanese plants that are used traditionally for neurological disorders as Alzheimer's disease, epilepsy and affective disorders as depression were tested for inhibition of acetylcholinesterase and affinity to the GABA (A)-benzodiazepine site and to the serotonin transporter. Ethyl acetate extracts of *Artemisia herba-alba* exhibited weak activity in the acetylcholinesterase assay. An ethanolic extract of *Artemisia herba-alba* had good affinity to the GABA (A)-benzodiazepine receptor site [116].

Dinatin and skrofulein, two flavones were extracted from *Artemisia herba-alba*, inhibited the binding of [methyl-3H]diazepam to rat brain membranes in vitro with IC_{50} of 1.3 and 23 µM, respectively. The GABA-ratios (the ratio of IC_{50} values in the absence/presence of GABA in the binding assay) were 1.1 and 1.2 for dinatin and skrofulein, respectively. Both flavones induced a slight increase in [³⁵S] TBPS binding. The data suggest that the flavones are antagonists or partial agonists of benzodiazepine receptors [117, 118].

6.10 Hypoglycemic activity

Artemisia herba-alba is a popular folk remedy, used in treatment of diabetes mellitus. For these reason many published papers studied its hypoglycemic activity [119, 120]; thus Twaij found that the aqueous extract of the aerial parts of A. herba-alba caused a significant fall in plasma glucose levels in both normoglycemic and the alloxanized rabbits [121]. Al-Waili reported that the water soluble extract of this species played a role in reducing elevated blood pressure and blood glucose levels, and it also had antiarrhythmic and anxiolytic actions [122, 123]. The same results are given in later study by Ibrahim [124]. Al-Yahya [125] reported that the ethanolic and chloroform extract of the whole plant had no effect on blood glucose levels. Al-Khazraji [126] studied the Artemisia herba-alba aerial parts and demonstrated that the aqueous extract of the roots does not contain hypoglycaemic agents, but the aqueous extract of the leaves was relatively more effective than the aqueous extract of the bark. He also found that the methanolic extract of the plant did not significantly change blood glucose levels of fasted rats. Al-Shamaony [127] gave a clear view that A. herba-alba extract prevented a significant elevation in the glycosylated haemoglobin in both diabetic rats and rabbits, fed daily for 2-4 weeks with A. herba-alba extract. Marrif [110] reported that the aqueous extract of A. herba-alba produced an initial hyperglycaemia which was followed by hypoglycaemia in normoglycaemic and alloxan-treated rabbits and mice. The extract also had no effect on the activity of alkaline phosphatase or concentrations of creatinine and urea in plasma. Also, Essway [128] concluded that the volatile oil of A. herba-alba (sage) have an antidiabetic effect after injection to rats. The study of the aqueous extract of A. herba-alba by Tastekin [129] came in agreement with Twaij [116] and Marrif [110] studies. Also, Bennani-Kabchi [130] reported that Artemisia herba-alba is a hypoglycemic, antihyperglycemic plant in diabetic sand rat, with hypolipemic action in case of severe obesity. A. herbaalba is one of the traditional plants used in Turkey, and the aqueous extract of A. herba-alba had been studied lately by Iriadam [131], who studied the effect of A. herba-alba and Teucrium polium effect on blood glucose levels. The results concluded that administration A. herba-alba might be useful in preventing hyperglycaemia by having insulin-like action and can significantly reduce the blood glucose in normoglycemic and hyperglycaemic rabbits but do not shows any action on creatinine and cholesterol in both normal and STZ-induced diabetic rabbits in acute diabetic experiment.

The aqueous extracts of *Artemisia herba-alba* has been tested on male albino rats. The results demonstrated that administration of *A. herba-alba* in a dose of 100 mg/kg of body weight for 60 days induced a very significant decrease in glucose level with a significant (p < 0.01) increase in total serum cholesterol level, triglycerides, phospholipids, serum aspartate aminotransferase (AST) and serum alanine aminotranserase (ALT) were found [132].

6.11 Cytotoxicity and gene induction

Essential oil (EO) extracted from *Artemisia herba-alba* was tested for its genotoxic effects using the yeast *Saccharomyces cerevisiae*. Clear cytotoxic effects were observed in the diploid yeast strain D7, with the cells being more sensitive to EO in exponential than in stationary growth phase. Clearly induced cytoplasmic peptide mutations took place indicating damage to mitochondrial DNA. However, no nuclear genetic events such as point mutations or mitotic intragenic or intergenic recombination were induced. The

capacity of EO to induce nuclear DNA damage-responsive genes was tested using suitable Lac-Z fusion strains for RNR3 and RAD51, which are genes involved in DNA metabolism and DNA repair, respectively. At equitoxic doses, the EO demonstrated significant gene induction, approximately the same as that caused by hydrogen peroxide, but much lower than that caused by methyl methanesulfonate (MMS). Essential oils affect mitochondrial structure and function and can stimulate the transcriptional expression of DNA damageresponsive genes. The induction of mitochondrial damage by EO appears to be closely linked to overall cellular cytotoxicity and appears to mask the occurrence of nuclear genetic events. EO-induced cytotoxicity involves oxidative stress, as is evident from the protection observed in the presence of ROS inhibitors such as glutathione, catalase or the iron-chelating agent deferoxamine [133]. In contentious study, investigated for their possible antigenotoxic effects in an eukaryotic cell system, the yeast Saccharomyces cerevisiae. The EO alone showed some cytotoxicity and cytoplasmic petite mutations, i.e. mitochondrial damage, but they were unable to induce nuclear genetic events. In combination with exposures to nuclear mutagens such as 254-nm UVC radiation, 8-methoxypsoralen (8-MOP) plus UVA radiation and methylmethane sulfonate (MMS), treatments with these EO produced a striking increase in the amount of cytoplasmic petite mutations but caused a significant reduction in revertants and mitotic gene convertants induced among survivors of the diploid tester strain D7. In a corresponding rho0 strain, the level of nuclear genetic events induced by the nuclear mutagens UVC and 8-MOP plus UVA resulted in the same reduced level as the combined treatments with the EO. This clearly suggests a close relationship between the enhancement of cytoplasmic petites (mitochondrial damage) in the presence of the EO and the reduction of nuclear genetic events induced by UVC or 8-MOP plus UVA. After MMS plus EO treatment, induction of these latter events was comparable at least per surviving fraction in wild type and rho0 cells, and apparently less dependent on cytoplasmic petite induction. Combined treatments with MMS and EO clearly triggered switching towards late apoptosis/necrosis indicating an involvement of this phenomenon in EO-induced cell killing and concomitant decreases in nuclear genetic events. After UVC and 8-MOP plus UVA plus EO treatments, little apoptosis and necrosis were observed. The antigenotoxic effects of the EO appeared to be predominantly linked to the induction of mitochondrial dysfunction. [134].

6.12 Reproductive toxic effects

Artemisia herba-alba toxic effect was investigated on the reproductive system after administration to female Sprague-Dawley rats weighting 250-300 g for two time periods 4 and 12 weeks. The effect of *A. herba-alba* exposure on fertility was assessed in terms of the number of pregnant rats, implantation sites, viable fetuses and resorption sites. Exposure to *A. herba-alba* for 4 weeks did not have much effect on fertility. Significant decrease in the relative ovarian weights and embryo weights in rats exposed to *A. herba-alba* were observed. Exposure to *A. herba-alba* for a 12 weeks resulted in a reduction in the percentage of pregnancies and in the number of implantation sites when compared with controls in both treatment periods. Rats receiving 12 weeks treatment showed an increase in ovarian weights and a decrease in the number of viable fetuses. These results indicate that long-term exposure of female rats to *A. herba-alba* causes adverse effects on the reproductive system and fertility. The results of the current study suggest that ingestion of *Artemisia herba-alba* by adult female rats causes adverse effects on fertility and reproduction [135].

Treated rats testicular cell population showed a decrease in number of spermatocytes and spermatids (p <0.01) when compared to controls. Serum hormonal assay indicated a decrease in testosterone and follicular stimulating hormone (FSH) levels in treated rats. A decrease in the number female rats impregnated by males receiving treatment was observed and demonstrated by a decrease in the implantation sites and number of viable fetuses (p < 0.01) [132].

6.13 Pesticidal activity

The potential of crude extracts from *Artemisia herba-alba* were evaluated for their ability to control *Tetranychus cinnabarinus* mites under controlled conditions. The assay revealed that the extract from *A*. *herba-alba* was not toxic to animal cells up to concentration of 100 μ g/mL. The mortality and repellency

potentials of the extract at a concentration of 50 μ g/mL caused only 58% and 71% mortality and repellency [136].

Artemisia herba-alba essential oil was tested against three sucking insect pests under laboratory and greenhouse conditions. These pests included *Bemisia tabaci* (Gennadius), *Aphis gossypii* (Glover) and *Thrips tabaci* (Lindman). The results showed that the LC50 of *A. herba-alba* was 0.042% for eggs and 0.074% for immature stages of *B. tabaci*. Also, the oil showed a high toxicity on *A. gossypii* with LC50 0.023 and 0.085%. *Artemisia herba-alba* was more toxic on *T. tabaci* and *A. gossypii* than *B. tabaci* in the laboratory test while *T. tabaci* was sensitive (LC50 0.011 and 0.038%). The oil was efficient for controlling tested insects on cucumber plants at greenhouses. This treatment caused 85.41, 83.57% reduction. in the population of *B. tabaci*, 90.44, 88.00% for *A. gossypii* and 87.45, 84.45% for *T. tabaci* [137, 138].

A study was designed to evaluate the effect of essential oils extracted from *Rosmarinus officinalis* and *Artemisia herba-alba* on *Acanthoscelides obtectus* (Coleoptera: Bruchidae) under laboratory conditions. The doses used were 1 to 5 μ L/30 g seed for the essential oil of each plant. The results show that the two tested essential oils were very toxic to adults of *A. obtectus*, and they also cause a significant reduction in fertility by bruchids. The *LD*₅₀, calculated after 48 hours of exposure, showed that the essential oil extracted from *Rosmarinus officinalis* was the more toxic to adults with an *LD*₅₀ = 0.59 μ L/30g seed, while the *LD*₅₀ = 1.69 μ L/30g seeds for *Artemisia herba-alba* [139]. larvicidal activity of *A. herba alba* against *Culex pipiens* mosquito and cotton leafworm *Spodoptera littoralis* (Biosd.) larvae, insecticidal activity against houseflies *Musca domestica* L. and rodenticidal activity against white mice *Mus musculus*, [111].

6.14 As Antibiotic resistant inhibitor

There are indications that some herbal materials can act as inhibitors of antibiotic resistance. A study was carried out to screen the effect of the herbal material in combination with antibiotics to identify systems that might be used advantageously to improve the efficiency of the antibiotics used. Preliminary results showed that saturated solutions of the plant extracts gave different levels of inhibition depending on the strain of microorganism. *Artemisia herba-alba*, for example, showed some inhibitory effects on the resistant strain and on standard strain of *S. aureus* [140].

6.15 Allelopathic effect

Plant-plant interference in semiarid Mediterranean environments may involve competition for resources, facilitation, or allelopathic phenomena in which toxic organic compounds produced by one plant are released into the environment. The potential allelopathic role of *Artemisia herba-alba* has been evaluated in order to explain the community pattern of the gypsum semiarid environments of central Spain. This pattern shows a sharp ecotone between a gypsophile spare shrubby community dominated by *Helianthemum squamatum*, which grows on slopes with gypsum surface crusts, and a nitrohalophilous community on the gypsum alluvial soils of piedmont dominated by *Artemisia herba-alba*. In order to explain the pattern, resource limitation was discarded because there were no significant differences in several soil parameters. Results confirm the inhibitory effect of aqueous extracts on the final germination percentage of scarified seeds of *Helianthemum squamatum* and also on the shape of the germination curves, which indicate delay of germination was strongly inhibited in soils obtained below the canopy of mature plants and retarted in other treatments [141].

6.16 Inhibition of the binding of 3H-benzo[a]pyrene

The aqueous and ethanolic plant extracts obtained from 10 Iraqi cultivated plant species were studied for their ability to inhibit aryl hydrocarbon hydroxylase (AHH) activity and [3H]benzo[a]pyrene ([3H]BP) binding to rat liver microsomal protein. None of the aqueous extracts tested showed any inhibitory effect. The ethanolic extracts from 8 species exhibited strong inhibitory effect on both AHH and [3H]BP binding to the microsomal protein. Ethanolic extracts from *Adiantum capillus-veneris* and *Salsola rosamarinus*

showed no effect. Nevertheless, ethanolic extracts from *Anchusa strigosa*, *Myrtus communis*, and *Artemisia herba-alba* were more effective than other plant extracts [142].

6.17 Cytoprotective properties of A. herba-alba

In a study by Gharzouli [143], investigated cytoprotective properties of Q. *ilex*, P. granatum and A. *herba-alba* aqueous extracts against ethanol-induced damage of the stomach. The gastroprotective effect of tannic acid and the aqueous extract of *Quercus ilex* L. root bark, *Punica granatum* L. fruit peel and *Artemisia herba-alba* Asso leaves were investigated in the rat against ethanol-induced damage. Tannic acid, Q. *ilex* and P. granatum extracts gave 100% precipitation of ovine haemoglobin *in vitro*, whereas A. *herba-alba* extract was devoid of any protein-binding property. Oral administration of these plant extracts or tannic acid induced a significant decrease in gastric lesions (47.7%– 76%). The observed protection was more pronounced when the test solution was given at the same time with ethanol, except for Q. *ilex* extract. The acid content of the stomach was significantly increased by P. granatum (368%) and A. *herba-alba* (251%) extracts prepared in ethanol. It is suggested that monomeric and polymeric polyphenols can strengthen the gastric mucosal barrier.

7. Effect of water stress on physiological and biochemical behavior of A. herba-alba.

Guenaoui aimed to study and investigate the response of *A. herba-alba* to water deficit stress. Plants were submitted for 36 days to 70 and 40 % CC and the reversibility of water deficit effect was assessed by analyzing the behaviour of water-stressed plants, once transferred to 100% CC treatment for 40 days. Several parameters known as indicators of plant status were monitored after drought treatment and subsequent re-watering. It was specially, about biomass production, plant water content, chlorophyll, sugars and praline contents and osmotic potential. Water drought led to a significant decrease of biomass accumulation, water content and chlorophyll content under both 70% and 40% CC. however, the intensity of water stress showed a positive correlation with sugars and praline accumulation. Well watered plants showed the higher biomass production but the lowest organic components unlike treated plants which showed the contrary evolution of these parameters. Drought stress, again, had a significant impact on the osmotic potential of plants, thereby it showed a significant decrease (P< 0.05) when irrigation was reduced to 70% CC or 40% CC. the stop of stress and the re-watering allowed the progressive return to the initial values and helped in the recovery of plants. Altogether, these data indicated that *A. herba-alba* has high aptitude permitting adaptation to unkind environmental conditions of the natural habitat [144].

8. Thermodynamic properties and moisture sorption isotherms of A. herba-alba

Sorption isotherms of *A. herba-alba* were determined at three temperatures (30, 40 and 50 °C) and in the range of water activity varying from 0.0572 to 0.898. The hysteresis effect was distinctly observed in the range of temperature tested. Five sorption models were used to fit the experimental data. The GAB equation was the best model describing the equilibrium moisture data for desorption. The modified Halsey equation was the most suitable model for describing adsorption isotherms. Thermodynamic properties such as differential enthalpy and entropy were determined from moisture sorption data, using the Clausius-Clapeyron equation. The experimental data showed that enthalpy-entropy compensation theory was applicable for the moisture sorption behaviour of *Artemisia herba-alba* [145].

9. Genetic polymorphism study of A. herba-alba

The morphological and chemical polymorphism observed for *A. herba-alba* may be due to a genetic variability [146]. To investigate this variability, Mohsen and Ali [1] studied the genetic polymorphism of

this plant on DNA level using inter simple sequence repeats (ISSR). The result of this study show that a total of 60 polymorphic loci were scored using our primers revealed no direct relation between morphological traits, geographic distance and genetic distance. Correlogram analysis showed a patchy distribution of the genetic variability of *A. herba-alba* accessions revealing the contribution of local ecological and geographic conditions on variability.

10. Microbial and heavy metals contamination

Alwakeel was conducted to evaluate the microbial contaminants and presence of toxic heavy metals on some herbal medicines. Twenty-seven samples (3 kg each) of well-known herbs and 5 kinds of henna available in herb markets around Riyadh, Saudi Arabia were collected for microbial and toxic metal contamination. *Artemisia herba-alba* was among the tested plants, which show the highest in calcium [147].

11. Inhibition corrosion of steel

Benabdellah tested *Artemisia* oil which is extracted from *Artemisia herba-alba* collected in Ain es-sefra-Algeria, as corrosion inhibitor of steel in 2 M H_3PO_4 using weight loss measurements, electrochem. polarization and EIS methods. The naturally oil reduces the corrosion rate. The inhibition efficiency was found to increase with oil content to attain 79% at 6 g/l. Ar acts as a cathodic inhibitor. The effect of temperature on the corrosion behavior of steel indicates that inhibition efficiency of the natural substance decreases with the rise of temp. The adsorption isotherm of natural product on the steel was determined [148]. Also, the Essential oil from *Artemisia herba-alba* tested as corrosion inhibitor of steel in 0.5 M H_2SO_4 using weight loss measurements and electrochem. polarization methods. Results gathered show that this natural oil reduced the corrosion rate by the cathodic action. Its inhibition efficiency attains the max. (74%) at 1 g/L. The inhibition efficiency of Arm oil increases with the rise of temp. The adsorption isotherm of natural product on the steel has been detd. *A. herba-alba* essential oil was obtained by hydrodistillation and its chemical oil compounds were investigated by capillary GC and GC/MS. The major components were chrysanthenone (30.6%) and camphor (24.4%) [149].

12. Biological effects to Artemisia herba-alba

12.1 G6PDH AND 6PGDH activities

Phenolic compounds were found to be phytotoxic constituents of many plant species extracts. Phenolic compounds caused seeds dormancy, inhibit seed germination and suppress photosynthesis in Artemisia herba-alba. Phenolic compounds are primarily synthesized through the pentose phosphate, and shikimate pathway. Glucose-6-phosphate dehydrogenase (G6PDH) and 6-phosphogluconate dehydrogenase (6PGDH) are the key enzymes of pentose phosphate pathway, used in the biosynthesis of phenolic compound. Al-Quadan and Al-Charchafchi studied the role played by the Phenolic compound esculetin in regulating pentose phosphate pathway activity in Artemisia herba-alba during seed germination. This is accomplished by studding G6PDH and 6PGDH activities in the presence and absence of esculetin. The result of these study showed that germination of Artemisia herba-alba seeds in the presence of esculetin showed lower G6PDH or 6PGDH activities compared to that germinated in esculetin absence; however, the same enzymes showed higher activities upon germination in the dark compared to that under light. Both enzymes were continuously increased from day one to day five during germination [150]. Al-Quadan, Ibrahim and Al-Charchafchi measured also the effect of Chlorogenic and caffeic acids on G6PDH and 6PGDH activites of A. herba-alba seeds during early stages of seed germination under conditions of light and dark. G6PDH and 6PGDH activities were inactivated in the presence of both phenolic acids and were stimulated upon germination in the dark. This study provide evidence or the variable needs of pentose phosphate pathway

activity, depending on concentrations of phenolic acids present and /or light conditions; moreover, a clear response of such germination condition was earlier expression of a second isoenzyme for G6PDH in the presence of 0.4 mM Chlorogenic or caffeic acids with apparently variable inhibitory pattern [151, 152].

Acknowledgment

The authors thank the late Prof. Ahmed A. Ahmed for his help and assistance.

References

- [1] H. Mohsen and F. Ali (2008). Study of genetic polymorphism of *Artemisia herba-alba* from Tunisia using ISSR markers, *African J. of Biotechnol.*, **7**(1), 44-50.
- [2] T. Efferth (2007). Antiplasmodial and antitumor activity of artemisinin-from bench to beside, *Planta Med.*, **73**, 299-309.
- [3] H. Wong and G. D. Brown (2002), Germacranolides from *Artemisia myriantha* and their conformation, *Phytochemistry*, **59**, 529-536.
- [4] J. H. Kwak, W. Y. Jang, O. P. Zee and R. K. Lee (1997). A new Coumarin-Monoterpene ether from Artemisia keiskeana, Planta Med., 63, 474-476.
- [5] R. X.Tan, H. Lu, J.-L. Wolfender, T. T. Yu, W. F. Zheng, L. S. Yang, Gafner and K. Hostettmann (1998). Mono-and sesquiterpenes and antifungal constituents from *Artemisia* species, *Planta Med.*, 65, 64-67.
- [6] O. Bergendorff and Sterner O. (1995). Spasmolytic flavonols from Artemisia *abrotanum*, *Planta Med.*, **61**, 370-371.
- [7] J. H. Kim, H-K. Kim, S. B. Jeon, K-H. Son, E. H. Kim, S. K. Kang, N-D. Sung and B-M. Kwon (2002). New sesquiterpene-monoterpene lactone, artemisolide, isolated from *Artemisia argyi, Tetrahedron lett.*, 43, 6205-6208.
- [8] S.-J. Lee, H-Y. Chung, C. G.-A. Maier, A. R. Wood, R. A. Dixon and T. J. Mabry, (1998). Estrogenic flavonoids from Artemisia vulgaris L, J. Agric. Food Chem., 46, 3325-3329.
- J. Hu, Q. Zhu, S. Bai and Z. Jia, (1996). New eudesmane sesquiterpene and other constituents from Artemisia mongolica, Planta Med., 62, 477-478.
- [10] M. N. Todorova, E. T. Tsankova, A. B. Trendailova and C. V. Gussev (1996). Sesquiterpene lactones with the uncommon rotundane skeleton from *Artemisia pontica* L, *Phytochemistry*, 41(2), 553-556.
- [11] R. Segal, I. Feuerstein and A. Danin (1987). Chemotypes of Artemisia herba-alba in Israel based on their sesquiterpene lactone and essential oil constitution. *Phytochemistry*, 15(4), 411-416.
- [12] A. Ziyyat, A. Legssyer, H. Mekhfi, A. Dassouli, M. Serhrouchni and W. Benjelloun (1997). Phytotherapy of hypertension and diabetes in oriental Morocco. J. of Ethnopharmacol., 58(1), 45-54.
- [13] A. Tahraoui, J. El-Hilaly, Z. H. Israili, B. Lyoussi (2007). Ethnopharmacological survey of plants used in the traditional treatment of hypertension and diabetes in south-eastern Morocco (Errachidia province), J. of Ethnopharmacol., 110 (1), 105-17.
- [14] N-A. Zeggwagh, O. Farid, J. B. Michel, M. Eddouks and I. Moulay (2008). Cardiovascular effect of Artemisia herba-alba aqueous extract in spontaneously hypertensive rats. Methods and findings in experimental and clinical pharmacology, 30(5), 375-81.
- [15] M. Laid, M-E. F. Hegazy and A. A. Ahmed (2008). Sesquiterpene lactones from Algerian Artemisia herbaalba. Phytochemistry lett., 1, 85-88.
- [16] J. Friedman, Z. Yaniv, A. Dafni and D. Palewitch (1986). A preliminary classification of the healing potential of medicinal plants, based on a rational analysis of an ethnopharmacological field survey among Bedouins in the Negev desert, Israel. *J. of Ethnopharmacol.*, **16**(2-3), 275-287.
- [17] F. Fenardji, M. Klur, Mrs. C. Fourlon and R. Ferrando (1974). White *artemisia*, (*Artemisia herba-alba*), *Revue d'Elevage et de Medecine Veterinaire des Pays Tropicaux*, **27**(2), 203-6.
- [18] A. Benmansur, S. A. Taleb-Bendiab, N. Mashev and G. Vasilev (1990). Studies on the chemical composition of *Artemisia (Artemisia herba-alba)*. *Bolgarskoi Akademii Nauk*, **43(8)**, 65-7.
- [19] A. Benmansour and S. A. taleb Bendiab (1998). Comparative investigation of proteins and amino acids in Artemisia herba-alba residues and Algerian date stones. Proposal to use them as additional feed for livestock. J. de la Societe Algerienne de Chimie, 8(1), 67-71.

- [20] L. Callegari and A. Rossi (1939). The active principles of Lybian Artemisia herba-alba and their pharmacological action. Archivio Italiano di Scienze Farmacologiche, 8, 145-60.
- [21] L. Callegari and A. Rossi (1940). The active principles of Lybian Artemisia herba-alba and their pharmacological action. Chimie et Industrie (Paris), 44, 321.
- [22] N. Feinbrun-Dothan (1978). *Flora Palaestina*, part **3**, 351-353, The Israel Academy of sciences and humanities, Jerusalem.
- [23] L. Boulos (2002), Flora of Egypt, vol. III. Al-Hadara publishing, Cairo, Egypt.
- [24] M. Torrell, M. Cerbah, S. Siljak-Yakovlev and J. Valles (2003). Molecular cytogenetics of the genus Artemisia (Asteraceae, Anthemideae): fluorochrome banding and fluorescence in situ hybridization. I. Subgenus Seriphidium and related taxa. Plant Systematics and Evolution, 239 (1-2),141-153.
- [25] J. Valles, M. Torrell, T. Garnatje, N. Garcia-Jacas, R. Vilatersana and A. Susanna (2003). The genus Artemisia and its allies: phylogeny of the subtribe Artemisiinae (Asteraceae, Anthemideae) based on nucleotide sequences of nuclear ribosomal DNA internal transcribed spacers (ITS). *Plant Biology (Stuttgart, Germany)*, 5(3), 274-284.
- [26] S. Salido, L. R. Valenzuela, J. Altarejos, M. Nogueras, A. Sanchez and E. Cano (2004). Composition and infraspecific variability of *Artemisia herba-alba* from southern Spain. *Biochem. Syst. Ecol.*, 32, 265-277.
- [27] C. E. Dunn, R. R. Brooks, J. Edmondson, M. Leblanc and R. D. Reeves (1996). Biogeochemical studies of metal-tolerant plants from southern Morocco. *Journal of Geochemical Exploration*, 56 (1), 13-22.
- [28] S. M. Khafagy, S. A. Gharbo and T. M. Sarg (1971). Phytochemical investigation of *Artemisia herba-alba*. *Planta med.*, **20**(1), 90-96.
- [29] R. Segal, S. Sokoloff, B. Haran, D. V. Zaitschek and Lichtenberg (1977). New sesquiterpene lactones from Artemisia herba-alba, Phytochemistry, 16, 1237-1241.
- [30] R. Segal, I. Feuerstein, H. Duddeck, M. Kaiser and A. Danin (1983). The sesquiterpene lactones from two populations of *Artemisia herba-alba*. *Phytochemistry*, 22(1), 129-131.
- [31] R. Segal, L. Eden, A. Danin, M. Kaiser and H. Duddeck (1984). Sesquiterpene lactones from a further population of *Artemisia Herba-alba*. *Phytochemistry*, **23**(12), 2954-2956.
- [32] R. Segal, L. Eden, A. Danin, M. Kaiser and H. Duddeck, (1985). Sesquiterpene lactones from Artemisia herba-alba. Phytochemistry, 24(6), 1381-1382.
- [33] R. Segal, I. Feuerstein and A. Danin (1987). Chemotypes of Artemisia herba-alba in Israel based on their sesquiterpene lactone and essential oil constitution. *Phytochemistry*, 15(4), 411-416.
- [34] S. E. Hull and O. Kennard (1978). 9β -Acetoxy-6(β H),11(β H)germacra-1β ,10α -epoxy,4(5)trans-ene-6,12olide, C17H24O5. Crystal Structure Communications, 7(1), 85-90.
- [35] J. D. Gomis, J. A. Marco, J. R. P. Llinares, J. S. Parareda, J. M. Sendra and E. Seoane (1979). Sesquiterpene lactones, waxes and volatile compounds from *Artemisia herba-alba* subspecies valentina. *Phytochemistry*, 18, 1523-1525.
- [36] J. A. Marco (1989). Sesquiterpene lactones from Artemisia herba-alba subsp. Herba-alba. Phytochemistry, 28(11), 3121-3126.
- [37] J. F. Sanz, G. Castellano and J. A. Marco (1990). Sesquiterpene lactones from Artemisia herba-alba. Phytochemistry, 29 (2), 541-545.
- [38] J. F. Sanz, E. Falco and J. A. Marco (1990). Further new sesquiterpene lactones from Artemisia herba-alba subsp. Valentine. J. Nat. Prod., 53(4), 940-945.
- [39] J. A. Marco, J. F. Sanz and E. Falco (1990). New oxygenated Eudesmanolides from Artemisia herba-alba. Tetrahedron, 46 (23), 7941-7950.
- [40] R. Bermudez, Y. Suarez (1912). Chemical analyses of forage plants of Spain. Langa y Compania, 94 pp.
- [41] J. F. Sanz and J. A. Marco (1991). New eudesmanolide related to Torrentin from Artemisia herba-alba subsp. Valentine. Planta med., 57, 74-76.
- [42] M. M. Gordon, D. V. Derveer and L. H. Zalkow (1981). New germacranolides from *Artemisia herba-alba*. X-RAY crystal structures of 3-8-dihydroxy-6-H,7-H,11-H-germacran-4(14), 9(10)-dien-6,12-olide and the corresponding 3-oxo-olide. *J. Nat. Prod.*, **44**(**4**), 432-440.
- [43] A. A. Ahmed, M. Abou-El-Ela, J. Jakupovic, A. A. Seif El-Din and N. Sabri (1990). Eudesmanolides and other constituents from *Artemisia herba-alba*. *Phytochemistry*, **29** (11), 3661-3663.
- [44] J. A. Marco, J. F. Sanz-Cervera and G. Ocete (1994). New germacranolides and Eudesmanolides from north African *Artemisia herba-alba*. J. Nat. Prod., **57**(7), 939-946.
- [45] D. Boriky, M. Berrada, M. Talbi, G. Keravis and F. Rouessac (1996). Eudesmanolides from Artemisia herbaalba. Phytochemistry, 43 (1), 309-311.
- [46] N. A. M. Saleh, S. I. El-Negoumy, M. F. Abd-Alla, M. M. Abou-Zaid, G. Dellamonica and J. Chopin (1985). Flavonoids glycosides of *Artemisia monosperma* and *A. herba-alba. Phytochemistry*, 24 (1), 201-203.

- [47] N. A. M. Saleh, S. I. El-Negoumy and M. M. Abou-Zaid (1987). Flavonoids of Artemisia judaica, A. monosperma and A. herba-alba. Phytochemistry. 26 (11), 3059-3064.
- [48] S. M. Salah and A. K. Jager (2005). Two flavonoids from Artemisia herba-alba Asso with in vitro GABA_Abenzodiazepine receptor activity. J. of Ethnopharmacol. 99, 145-146.
- [49] R. Segal, D. Cohen, S. Sokoloff and D. V. Zaitschek (1973). New flavone from Artemisia herba-alba. Lloydia , 36(1), 103-5.
- [50] F. Mouhajir, J. A. Pedersen, M. Rejdali and S. G. H. N. Touer (2001). Phenolics in Moroccan medicinal plant species as studied by electron spin resonance spectroscopy. Can Pharmaceutical Biology, **39** (5), 391-398.
- [51] T-H. Kim, H. Ito, T. Hatano, S. Taniguchi, S. Khennouf qnd T. Yoshida (2004). Chemical constituents of *Artemisia herba-alba* Asso., *Nat. Med.* **58**(4), 165.
- [52] P. I. Sanchez and G. J. Delgado (1978). Organic principles in east Spanish plants. III. Separation and identification of the components of an *Artemisia Herba-alba* Valentina wax by gas-liquid chromatography. Grasas y Aceites (Sevilla, Spain), 29(5), 323-7.
- [53] B. M. Lawrence (1994). Progress in essential oils, Perfume. & Flavor., 19, 83-84, 86, 88-92, 94-95.
- [54] B. M. Lawrence (1989). Progress in essential oils, perfume. & flavor., 14(3), 71-6, 78-80.
- [55] B. M. Lawrence (1981). Progress in essential oils, perfume. & flavor., **6**(1), 37-8, 43-6.
- [56] S. Lemberg (1982). Armoise- Artemisia herba alba. Perfume. & flavor., 7, 58-60, 62-63.
- [57] J. Yashphe, I. Feuerstein, S. Barel and R. Segal (1987). The antibacterial and antispasmodic activity of Artemisia herba-alba Asso. II. Examination of essential oils from various chemotypes. Int. J. Crude-Drug-Res. 25 (2), 89-96.
- [58] Z. Fleisher, A. Fleisher and R. B. Nachbar (2002). Chemovariation of *Artemisia herba-alba* Asso. Aromatic plants of the Holy land and the Sinai. Part XVI. J. Essen. Oil Res., 14, 156-160.
- [59] U. Ravid, E. Putievsky and I. Katzir (1996). Stereochemical analysis of borneol in essential oils using permethylated β -cyclodextrin as a chiral stationary phase. *Flavour and Fragrance J.*, **11**(3), 191-195.
- [60] R. Segal, A. Breuer and I. Feuerstein (1980). Irregular monoterpene alcohols from *Artemisia herba-alba*. *Phytochemistry*, **19**, 2761-2762.
- [61] I. Feuerstein, D. Mueller, K. Hobert, A. Danin and R. Segal (1986). Constitution of essential oils from Artemisia herba-alba populations of Israel and Sinai. Phytochemistry, 25, 2343-2347.
- [62] A. M. El-Sayed and A. A. Seida (1990). Comparative study of the major constituents of the essential oils of wild and cultivated Egyptian Artemisia herba-alba with those of plants produced abroad. Bull. Fac. Pharm. (Cairo Univ.), 28, 57-58.
- [63] M. Hurabielle, M. Malsot and M. Paris (1981). Chemical study of two oils from wormwood; *Artemisia herbaalba* Asso and *Artemisia vulgaris* Linnaeus of chemicotaxonomic interest. *Riv. Ital.* EPPOS, **63**, 296-299.
- [64] M. Hudaib, T. Aburjai (2006). Composition of the essential oil from *Artemisia herba-alba* growing in Jordan, *J. Essen. Oil Res.*, **18**, 301-304.
- [65] A. Cohen, J. P. Lavergne, A. Leblanc (1972). Constituents of *Artemisia herba-alba* (typica). I. essential oils, *Bulletin de la société des sciences naturelles et physiques du Maroc.* **52(1-2)**, 1-19.
- [66] A. Ouyahya, R. Negre, J. Viano, Y. F. Lozano and E. M. Gaydou (1990). Essential oils from Moroccan Artemisia negrei, A. mesatlantica and A. herba-alba. Lebensm Wiss. Technol., 23, 528-530.
- [67] B. Benjilali and H. Richard (1980). Study of some populations of Artemisia herba-alba from Morocco. Riv. Ital. Essen., 62, 69-74.
- [68] B. Benjilali, J. Sarris and H. Richard (1982). New chemotypes of Artemisia herba-alba, Sci. Aliments, 2, 515-527.
- [69] A. Lamiri, A. Belanger, M. Berrada, S. Zrira, B. Benjilali (1997). Chemical polymorphism of Artemisia herba-alba Asso from Morocco (in French) Rabat: Morocco, 69-79.
- [70] A. Lamiri, A. Belanger, M. Berrada, M. M. Ismaili-Alaoui and B. Benjilali (1997). Origin of chemical polymorphism of Moroccan *Artemisia herba-alba* Asso (in French); *Rabat: Morocco*, 81-92.
- [71] I. Feuerstein, A. Danin, R. Segal (1988). Constitution of the essential oil from an Artemisia herba-alba population of Spain, *Phytochemistry*, **27**, 433-434.
- [72] E. Alinari (1924). The probable presence of thujone in the essential oil of Artemisia herba-alba Asso., Annali di chimica Applicata, 14, 109-12.
- [73] Fesneau, Maxime (1950). Spanish essential oils. Industries de la parfumerie, 5, 132-3.
- [74] F. Roure-Bertrand (1920). Essential oils, Sci. Ind. Bull. (Grasse). April (Chem. Abst. 1920, 14, 3752).

- [75] A. Villar, M. L. Caduch and M. C. Zafra-Polo (1983). Essential oils from different Artemisia species, Ars. Pharm. Aceutica, 24(2), 149-59.
- [76] A. Akrout (2004). Essential oil study of some pastoral plants from Matmata (south Tunisia)(in French). Cah. Options Med., 62, 289-292.
- [77] A. Neffati, I. Skandrani, M. B. Sghaier, I. Bouhlel, S. Kilani, K. Ghedira, M. Nefati, I. Cherif, M. Hammani and L. Chekir-Ghedira (2008). Chemical composition, mutagenic and antimutagenic activities of essential oils from (Tunisian) *Artemisia campestris* and *Artemisia herba-alba*. J. Essen. Oil Res., 20, 471-477.
- [78] M. Haouari and A. Ferchichi (2009). Essential oil composition of Artemisia herba-alba from southern Tunisia, Molecules, 14, 1585-1594.
- [79] G. Vernin and C. Parkanyi (2001). GC/MS analysis of Artemisia herba-alba Asso. from Algeria, Nonpolar and polar extracts. *Riv. Ital.* EPPOS, 32, 3-16.
- [80] G. Vernin, O. Merad, G. M. Vernin, R. M. Zamkotsian and C. D. Parkanyi (1995). GC/MS analysis of Artemisia herba-alba Asso essential oils from Algeria. Develop. in food Sci., 37A, 147-205.
- [81] G. Vernin and L. O. Merad (1994). Mass spectra and Kovats indexes of some new *cis*-chrysanthenyl esters found in the essential oil of *Artemisia herba-alba* from Algeria. J. Essen. Oil Res., 6, 437-338.
- [82] N. Dahmani-Hamzani and A. Baaliouamer (2005). Chemical composition of the Algerian essential oil of Artemisia herba-alba native to Dejelfa. Riv. Ital. EPPOS, 40, 7-13.
- [83] G. Vernin, O. Merad, R. M. Zamkotsian, G. M. F. Vernin and C. Parkanyi (1998). Book of Abstracts, 215th ACS National Meeting.
- [84] C. Boutekedjiret, S. Charchari, R. Belabbes, J. M. Bessiere (1992). Chemical composition of the essential oil of Artemisia herba-alba. Riv. Ital EPPOS, 3(8), 39-42.
- [85] S. Charchari and R. Belabbes (1991). Infrared spectroscopic determination of camphor and thujone in the essential oil and concrete of *Artemisia herba-alba* Asso. *Riv. Ital.* EPPOS, **2**(3), 13-16.
- [86] G. Emilien (1904). Essential oil of Artemisia herba-alba of Algeria. Bulletin de la Société chimique de France, **31(iii)**, 694-7.
- [87] E. Alinari (1922). Ethereal oil from Artemisia herba-alba, Riv. It. delle essenze e profumi, 4, 75-77.
- [88] S. Salido, J. Altarejos, M. Nogueras and A. Sanchez (2001). Chemical composition of essential oil of Artemisia herba-alba Asso. ssp. valentina (Lam.) Marc. J. Essen. Oil Res., 13, 221-224.
- [89] T. Dob, T. Benabdelkader (2006). Chemical composition of the essential oil of Artemisia herba-alba Asso grown in Algeria. J. Essen. Oil Res., 18, 685-690.
- [90] B. M. Lawrence (1993). Armoise oil, in: Essential oils 1988-1991. edit., *Natural Flavor and Fragrance materials "perfumer & Flavorist"*, p. 52-54, Allures publ. corp., carol stream, IL.
- [91] B. M. Lawrence (1995). Armoise oil. In: Essential oils 1982-1994 Edit., natural Flavor and Fragrance materials "Perfumer & Flavorist", p. 179-180, Allured Publ. Corp., carol stream, IL.
- [92] A. Maugini (1930). Some plants of Cyrenaica. Agr. coloniale (Florence), 24, 74-77, 127-131, 189-191.
- [93] A. Djeridane, M. Yousfi, B. Nadjemi, D. Boutassouna, P. Stocker and N. Vidal (2006). Antioxidant activity of some Algerian medicinal plants extracts containing phenolic compounds. *Food Chemistry*, 97, 654-660.
- [94] A. Amr (1995). Antioxidative role of some aromatic herbs in refrigerated ground beef patties. *Pure and Applied Sciences*, **22B(6)**, 1475-1487.
- [95] A. H. Al-Mustafa and O. Y. Al-Thunibat (2008). Antioxidant activity of some Jordanian medicinal plants used traditionally for treatment of diabetes. *Pak. J. of biol. sci.: PJBS*, **11(3)**, 351-8.
- [96] Z. B. Abid, M. Feki, A. Hédhili and M. H. Hamdaoui (2007). Artemisia herba-alba Asso (Asteraceae) has equivalent effects to green and black tea decoctions on antioxidant processes and some metabolic parameters in rats. Ann. Nutr. Metab., 51, 216-222.
- [97] S. M. Suboh, Y. Y. Bilto and T. A. Aburjai (2004). Protective effects of selected medicinal plants against protein degradation, lipid peroxidation and deformability loss of oxidatively stressed human erythrocytes. *Phytother. Res.*, 18(4), 280-4.
- [98] A-K. J. Sallal and A. Alkofahi (1996). Inhibition of the hemolytic activities of snake and scorpion venoms in vitro with plant extracts. *Biomedical Lett.*, 53(212), 211-215.
- [99] M. A. Saleh, M. H. Belal and G. EL-Baroty (2006). Fungicidal activity of Artemisia herba-alba Asso (Asteraceae). J. of envir. sci. and health. Part B. Pesticides, food contaminants, and agricultural wastes, 41(3), 237-244.
- [100] C. Bouchra, A. Mohamed, I. H. Mina and M. Hmamouchi (2003). Antifungal activity of essential oils from several medicinal plants against four postharvest citrus pathogens. *Phytopathologia Mediterranea*, 42(3), 251-256.

- [101] M. Khaddor, A. Lamarti, A. Tantaoui-Elaraki, M. Ezziyyani, M. Candela Castillo, A. Badoc (2006). Antifungal activity of three essential oils on growth and toxigenesis of Penicillium aurantiogriseum and Penicillium viridicatum. J. of Essen. Oil Res., 18(5), 586-589.
- [102] A. Tantaoui-Elaraki, H. Ferhout and A. Errifi (1993). Inhibitation of the fungal asexual reproduction stages by three Moroccan essential oils, J. Essen. Oil Res., 5(5), 535-545.
- [103] A. Tantaoui-Elaraki and A. Errifi (1994). Antifungal activity of essential oils when associated with sodium chloride or fatty acids. *Grasas y Aceites* (Seville), **45(6)**, 363-369.
- [104] G. Roger, H. Youcef and K. Jacques (2008). Compositions and antifungal activities of essential oils of some Algerian aromatic plants. *Fitoterapia*, **79(3)**, 199-203.
- [105] N. M. A. El-Shayeb and S. S. Mabrouk (1984). Utilization of some edible and medicinal plants to inhibit aflatoxin formation. *Nutrition Reports International*, **29(2)**, 273-82.
- [106] S. Charchari, A. Dahoun, F. Bachi and A. Benslimani (1996). Antimicrobial activity in vitro of essential oils of Artemisia herba-alba Asso and Artemisia judaica L. from Algeria. Riv. Ital. EPPOS, 18, 3-6.
- [107] L. Al-Banna, R. M. Darwish and T. Aburjai (2003). Effect of plant extracts and essential oils on root-knot nematode. *Phytopathologia Mediterranea*, 42(2), 123-128.
- [108] J. Yashphe, R. Segal, A. Breuer and G. Erdreich-Naftali (1979). Antibacterial activity of Artemisia herbaalba. J. of Pharma. Sci. 68(7), 924-925.
- [109] A. Sherif, R. G. Hall and M El-Amamy (1987). Drugs, insecticides and other agents from Artemisia. Medical Hypotheses, 23, 187-193.
- [110] H. Marrif, B. H. Ali and K. M. Hassan (1995). Some pharmacological studies on *Artemisia herba-alba* (Asso.) in rabbits and mice. *J. of Ethnopharmacol.*. **49**, 51-55.
- [111] M. S. Hifnawy, O. A. Rashwan and M. A. Rabeh (2001). Comparative chemical and biological investigations of certain essential oils belonging to families Asteraceae, Lamiaceae and Graminae. *Bulletin of the Faculty of Pharmacy* (Cairo University), **39**(2), 35-53.
- [112] W. Al-Momani, E. Abu-Basha, S. Janakat, R. A. J. Nicholas and R. D. Ayling (2007). In vitro antimycoplasmal activity of six Jordanian medicinal plants against three Mycoplasma species. *Trop. Anim. Health Prod.*, 39, 515-519.
- [113] U. E. Idris, S. E. Adam and G. Tartour (1982). The anthelmintic efficacy of *Artemisia herba-alba* against Haemonchus contortus infection in goats. *National Institute of Animal Health quarterly*, **22(3)**, 138-43.
- [114] N. S. Al-Waili (1988). Artemisia herba-alba extract for treating Enterobius vermicularis infection. Transactions of the Royal Society of Tropical Medicine and Hygiene, 82, 626.
- [115] S. Hatimi, M. Boudouma, M. Bichichi, N. Chaib and N. G. Idrissi (2001). Evaluation in vitro de l'activité antileishmanienne d'Artemisia herba-alba Asso. Thérapeutique, Bulletin de la Société de pathologie exotique, 94(1), 29-31.
- [116] S. M. Salah, A. K. Jager (2005). Screening of traditionally used Lebanese herbs for neurological activities. J. of Ethnopharmacol., 97(1), 145-9.
- [117] X. L. Shen, M. Nielsen, M. R. Witt, O. Sterner, O. Bergendorff and M. Khayyal (1994). Inhibition of [methyl-3H]diazepam binding to rat brain membranes in vitro by dinatin and skrofulein. *Acta pharmacologica Sinica*, 15(5), 385-8.
- [118] X. Shen, M. R. Witt, M. Nielsen, O. Bergendorff, O. Sterner, M. T. Khayyal, T. S. El-Alfy and H. El-Gohary (1993). Flavone derivatives isolated from *Artemisia herba-alba* have affinity to brain benzodiazepine and α 1adrenergic receptors in vitro. *Zhongguo Yaolixue Yu Dulixue Zazhi*, 7(4), 305-6.
- [119] H. Jouad, M. Haloui, H. Rhiouani, J. El-Hilaly and M. Eddouks (2001). Ethnobotenical survey of medicinal plants used or the treatment of diabetes, cardiac and renal diseases in the North centre region of Morocco (Fez-Boulemane). J. of Ethnopharmacol.. 77, 175-182.
- [120] M. Eddouks, M. Maghrani, A. Lemhadri, M.-L. Ouahidi and H. Jouad (2002). Ethnopharmacological survey of medicinal plants used for the treatment of *Diabetes mellitus*, hypertension and cardiac diseases in the Southease region of Morocco (Tafilalet). J. of Ethnopharmacol.. 82, 97-103.
- [121] H. A. A. Twaij and A. A. Al-Badr (1988). Hypoglycemic activity of Artemisia herba-alba. J. of Ethnopharmacol., 24, 123-126.
- [122] N. S. Al-waili (1986). Treatment of diabetes mellitus by Artemisia herba-alba extract: preliminary study. Clinic. and Experim. Pharmacol. and Physiol., 13, 569-573.
- [123] N. S. Al-Waili (1988). Artemisia herba-alba and diabetes mellitus. Clinical and Experim. Pharmacol. and Physiol.. 15(6), 497.

- [124] I. G. Ibrahim, B. El-Salkh, N. Shawki, S. M. Abou El-Fotouh and H. M. Abou El-Fotouh (2002). Comparative study on the effect of gliclazide and two antidiabetic plants used in folk medicine on albino rat's fetuses. *Egyptian Journal of Hospital Medicine*, 6, 80-98.
- [125] M. A. Al-Yahya, M. Tariq, I. A. Al-Meshal, S. J. Mossa and A. A. Al-Badr (1986). Artemisia herba-alba, Saudi plants: A phytochemical and biological Approach. College of pharmacy, King Saud University, Riyadh, pp. 44-49.
- [126] S. M. Al-Khazraji, L. A. Al-Shamaony and H. A. A. Twaij (1993). Hypoglycaemic effect of Artemisia herbaalba. I. effect of different parts and influence of the solvent on hypoglycaemic activity. J. of Ethnopharma., 40, 163-166.
- [127] L. A. Al-Shamaony, S. M. Al-Khazraji and H. A. A. Twaij (1994). Hypoglycaemic effect of Artemisia herbaalba. II. Effect of a valuable extract on some blood parameters in diabetic animals. J. of Ethnopharma.. 43, 167-171.
- [128] G. S. Essway, H. M. Sobbhy and H. A. El-Banna (1995). The hypoglycemic effect of volatile oil of some Egyptian plants. *Veterinary Medical Journal Giza*, 43(2), 167-172.
- [129] D. Tastekin, M. Atasever, G. Adigüzel, M. Keles and A. Tastekin (2006). Hypoglycaemic effect of Artemisia herba-alba in experimental hyperglycaemic rats. Bull. Vet. Inst. Pulawy. 50, 235-238.
- [130] N. Bennani-Kabchi, Y. Cherrah, H. Fdhil and G. Marquie (1995). Therapeutic effect of *Artemisia herba-alba* on lipidic and carbohydrate metabolism in diabetic sand rat (Psammomys obesus). *Pharmaco. Res.*, **31**, 381.
- [131] M. Iriadam, D. Musa, H. Gümühan and F. Baba (2006). Effects of two Turkish medicinal plants Artemisia herba-alba and Teucrium polium on blood glucose levels and other biochemical parameters in rabbits. J. of Cell and Molecular Biol.. 5, 19-24.
- [132] M. H. Khataibeh and H. Daradka (2007). Antiandrogenic activity of *Artemisia herba-alba* in male rats, with emphasis on biochemical parameters. *Asian J. of chemistry*, **19** (4), 2595-2602.
- [133] F. Bakkali, S. Averbeck, D. Averbeck, A. Zhiri and M. Idaomar (2005). Cytotoxicity and gene induction by some essential oils in the yeast Saccharomyces cerevisiae. *Mutation research*, 585 (1-2), 1-13.
- [134] F. Bakkali, S. Averbeck, D. Averbeck, A. Zhiri, D. Baudoux and M. Idaomar (2006). Antigenotoxic effects of three essential oils in diploid yeast (Saccharomyces cerevisiae) after treatments with UVC radiation, 8-MOP plus UVA and MMS. *Mutation research*, 606 (1-2), 27-38.
- [135] M. M. Almasad, W. Sh. Qazan and H. Daradka (2007). Reproductive toxic effects of Artemisia herba-alba ingestion in female Spague-dawley rats. Pak. J. of Bio. Sci. 10 (18), 3158-3161.
- [136] H. Azaizeh, M. Kobaisy, S. Dakwar, B. Saad, I. Shaqir and O. Said (2007). Botanical pesticides as a source of safe bioacaricides for the control of Tetranychus cinnabarinus. *Acta Phytopathologica et Entomologica Hungarica*, **42**(1), 143-152.
- [137] M. M. Soliman (2006). Phytochemical and toxicological studies of Artemisia L. (Compositae) essential oil against some insect pests. Acta Phytopathologica et Entomologica Hungarica, 41(3-4), 395-406.
- [138] M. M. Soliman (2007). Phytochemical and toxicological studies of Artemisia L. (Compositae) essential oil against some insect pests. Archives of Phytopathology and Plant Protection, 40 (2), 128-138.
- [139] Z. B. Tani, F. Hassani and M. A. Khelil (2008). Bioefficacy of essential oils extracted from the leaves of *Rosmarinus oficinalis* and *Artemisia herba-alba* towards the Bruche bean Acanthoscelides obtectus (coleoptra: Bruchidae). J. of Pure & Applied Microbiol. 2 (1), 165-170.
- [140] R. M. Darwish, T. Aburjai, S. Al-Khalil and A. Mahafzah (2002). Screening of antibiotic resistant inhibitors from local plant materials against two different strains of Staphylococcus aureus. J. of Ethnopharma., 79, 359-364.
- [141] A. Escudero, M. J. Albert, J. M. Pita and F. Pérez-Garcia (2000). Inhibitory effects of Artemisia herba-alba on the germination of the gypsophyte Helianthemum squamatum. Plant Ecology, 148, 71-80.
- [142] A. H. Alwan, K. A. S. Al-Gaillany and A. Naji (1989). Inhibition of the binding of 3H-benzo[a]pyrene to rat liver microsomal protein by plant extracts. *International J. of Crude Drug Res.*, **27**(1), 33-7.
- [143] K. Gharzouli, S. Khennouf, S. Amira and A. Gharzouli (1999). Effects of aqueous extracts from *Quercus ilex* L. root bark, *Punica granatum* L. fruit peel and *Artemisia herba-alba* Asso leaves on ethanol-induced gastric damage in rats. *Phytother. Res.*, 13, 42-45.
- [144] C. Guenaoui, M. Gorai, S. Smiti and M. Neffati (2008). Biochemical and physiological changes in *Artemisia herba-alba* plants under water stress conditions. *Middle-East Journal of Scientific Research*, **3** (**3**), 156-163.
- [145] A. Lamharrar, A. Idlimam and M. Kouhila (2007). Thermodynamic properties and moisture sorption isotherms of *Artemisia herba-alba*. *Revue des Energies Renouvelables*. **10** (3), 311-320.
- [146] M. Torrell and J. Valles (2001). Genome size in 21 Artemisia L. species (Asteraceae, Anthemideae): Systematic, evolutionary, and ecological implications. Genome, 44(2), 231-238.

- [147] S. S. Alwakeel (2008). Microbial and heavy metals contamination of herbal medicines. *Res. J. of Microbiol.*, 3 (12), 683-691.
- [148] M. Benabdellah, M. Benkaddour, B. Hammouti, M. Bendahhou and A. Aouniti (2006). Inhibition of steel corrosion in 2M H3PO4 by artemisia oil. *Applied Surface Science*, 252 (18), 6212-6217.
- [149] O. Ouachikh, A. Bouyanzer, M. Bouklah, J.-M Desjobert, J. Costa, B. Hammouti and L. Majidi (2009). Application of essential oil of *Artemisia herba-alba* as green corrosion inhibitor for steel in 0.5 M H2SO4. *Surface Review and Letters*, 6(1), 49-54.
- [150] F. Al-Quadan and F. M. R. Al-Charchafchi (2006). G6PDH and 6PGDH activities of *Artemisia herba-alba* seeds and seedlings during germination in presence of Esculetin. *Pak. J. of Biol. Sci.*. **9** (1), 210-213.
- [151] F. M. R. Al-Charchafchi and F. Al-Quadan (2006). Effect of chlorogenic acid on germination and seedling growth and on the enzymes activity extracted from *Artemisia herba alba* ASSO. Part I: germination and seedling growth. *Pure Sciences*. 33(2), 168-175.
- [152] F. Al-Quadan, A. Ibrahim and F. M. R. Al-Charchafchi (2008). Effect of Chlorogenic and caffeic acids on activities and isoenzymes of G6PDH and 6PGDH of *Artemisia herba-alba* seed Germinated for one and three days in light and dark. *Jordan J. of Biol. Sci..* 1(2), 85-88.



© 2010 Reproduction is free for scientific studies