

## Medicinal Plants Alkaloids, As a Promising Therapeutics- A Review (Part 1)

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### ABSTRACT:

Natural products have gained popularity worldwide for promoting healthcare, as well as disease prevention. Medical use of alkaloid-containing plants has a long history and many alkaloids are still used in medicine nowadays. Alkaloids possessed wide range of pharmacological and therapeutic effects. The current manuscript discussed the plant contents of alkaloids for medical, pharmaceutical, synthetic, and many other useful properties.

**KEYWORDS:** Medicinal plants, alkaloid, pharmacology, therapeutic

### I. INTRODUCTION:

Alkaloids are a huge group of naturally occurring organic compounds which contain nitrogen atom or atoms (amino or amido in some cases) in their structures. These nitrogen atoms cause alkalinity of these compounds. Alkaloids have a wide range of pharmacological effects included ajmaline (antiarrhythmic), colchicin (antigout), emetine (antiprotozoal agent, emesis), ergot alkaloids (vasoconstriction, hallucinogenic, uterotonic), glaucine (antitussive), morphine (analgesic), codeine (analgesic, antitussive), nicotine (nicotinic acetylcholine receptor agonist), physostigmine (acetylcholinesterase inhibitor), quinidine (antiarrhythmic), quinine (antipyretic, antimalarial), reserpine (antihypertensive), tubocurarin (muscle relaxant), vinblastine, vincristine (antitumor), vincamine (vasodilating, antihypertensive), yohimbine (stimulant, aphrodisiac), and many other therapeutic effects<sup>(1-3)</sup>. The current review was designed to highlight the plant contents of alkaloids for medical, pharmaceutical, synthetic, and many other useful properties.

#### Medicinal plants contain alkaloids

##### *Adiantum capillus-veneris*

The Soxhlet extraction of *Adiantum capillus-veneris* showed that it contained 0.53% alkaloids<sup>(4-5)</sup>.

##### *Ailanthus altissima*

Alkaloidal glycosides (canthin-6-one, 1-methoxycanthin-6-one, 5-methoxycanthin-6-one, and canthin-6-one-3-N-oxide), were isolated from the root bark of *Ailanthus altissima*<sup>(6-7)</sup> while, the leaves contained the alkaloid linuthine<sup>(8-11)</sup>.

##### *Ammi majus*

*Ammi majus* seeds contained 0.73 g ergot alkaloids (expressed as ergonovine) per 100g<sup>(12-14)</sup>.

##### *Anagallis arvensis*

The seeds of *Anagallis arvensis* contained 2.21 % alkaloids<sup>(15-16)</sup>.

##### *Anagyris foetida*

*Anagyris foetida* contained many alkaloids, included, anagryne, baptifoline, isorhamnetin, and syringin 4-O-beta-D-glucopyranoside. The crude alkaloid can be separated into cytisine, and anagryne, the latter is not obtainable in crystalline form; it forms a brittle, resinous-like mass easily reducible to a yellowish powder, but rapidly absorbs moisture and becomes sticky. It volatilizes at 245° C. (473° F.). It forms, however, a crystalline hydrobromide<sup>(17-19)</sup>.

##### *Anachus strigosa*

Highest total concentration of pyrrolizidine alkaloids was detected in the leaves of *Anachus strigosa* (23.63 mg/g of dried part), followed by the flowers (19.77 mg/g), and finally by the roots (1.80 mg/g). These pyrrolizidine alkaloids included 7,7'-bis-(4-hydroxy-3,5-dimethoxyphenyl)-8,8'-dihydroxymethyl tetrahydrofuran 4'-O-β-D-glucopyranoside, rosmarinic acid, caffeic acid, tormentic acid 28-O-β-D-glucopyranoside, euscaphic acid 28-O-β-D-glucopyranoside, euscaphic acid, and allantoin. However Braca et al isolated new six pyrrolizidine alkaloids included a new carboxylic acid, a new phenolic and a new oleanane glycoside<sup>(20-22)</sup>.

***Andrachne aspera***

The aerial parts of *Andrachne aspera* was shown to contain alkaloids, andrachamine, andrachcine, andrachcinine, andrachcinidine, (+)-allosedridine, (-)-8-epi-8-ethylnorlobelol and (-)-8-epihalosaline<sup>(23-24)</sup>. The aerial parts of *Andrachne aspera* is also contained many piperidine alkaloids, these included aspertin-A, aspertin-B, aspertin-C and aspertin-D<sup>(8)</sup>. Terpenes isolated from the aerial parts are lupeol acetate,  $\alpha$ -amyrin,  $\beta$ -amyrin,  $\alpha$ -taraxerol, stigmasterol,  $\beta$ -stigmasterol, lupeol, oleanolic acid and germanicol<sup>(25-27)</sup>.

***Antirrhinum majus***

*Antirrhinum majus* contained four tertiary alkaloids, one of which has been identified as 4-methyl-2,6-naphthyridine. A water-soluble base has been identified as choline<sup>(28-29)</sup>.

***Arachis hypogaea***

Two indole alkaloids were isolated from water-soluble fraction of peanut (*Arachis hypogaea*) skins. The alkaloids were identified as 2-methoxyl-3-(3-indolyl)-propionic acid and 2-hydroxyl-3-[3-(1-N-methyl)-indolyl]-propionic acid<sup>(30-31)</sup>.

***Aristolochia maurorum***

Three atypical alkaloids (aristolochic acid I, aristolochic acid II and aristolochic acid IIIa), have been isolated from *Aristolochia maurorum*<sup>(32)</sup>.

***Arundo donax***

*Arundo donax* contained alkaloids including tryptamine, bufotenidine, gramine and arundamine<sup>(33-35)</sup>.

***Astragalus tribuloides***

Indolizidine alkaloids, were isolated from *Astragalus tribuloides*<sup>(36-37)</sup>.

***Bacopa monniera***

*Bacopa monniera* contained alkaloid brahmine, nicotine, and herpestine<sup>(38-40)</sup>.

***Caccinia glauca***

An alkaloid, a diester of retronecine and benzoic acid was isolated from flowers of *Caccinia glauca*<sup>(41)</sup>.

***Capparis spinosa***

Stachydrine (a pyridine alkaloid), and cadabicine (a 24-membered polyamine lactam alkaloid) were isolated from *Capparis spinosa*<sup>(42-45)</sup>.

***Capsicum species***

Alkaloid compounds called capsaicinoids were isolated from the fruits of *Capsicum species*<sup>(46-49)</sup>.

***Casuarina equisetifolia***

Casuarine- a very highly oxygenated pyrrolizidine alkaloid was extracted from *Casuarina equisetifolia*<sup>(50-51)</sup>.

***Centaurea cyanus***

Analysis of a methanol extract of the seeds of *Centaurea cyanus* gave four indole alkaloids: moschamine, cis-moschamine, centcyamine and cis-centcyamine<sup>(52-53)</sup>.

***Chenopodium album***

Analysis of the leaves of four *Chenopodium album* cultivars showed that they contained alkaloids 1-27-1.53 mg/100g. Chenoalbicin, a novel cinnamic acid amide alkaloid was isolated from *Chenopodium album*<sup>(54-57)</sup>.

***Citrus Species***

*Citrus aurantifolia*, *Citrus limonum* and *Citrus sinensis* fruits contained: alkaloids: 0.33  $\pm$  0.11, 0.54  $\pm$  0.20, 0.62  $\pm$  0.10 mg/ 100g dry weight respectively<sup>(58-60)</sup>.

***Colchicum Species***

Nine alkaloids (3-demethylcolchicine, 2-demethylcolchicine, colchifoline, N-deacetyl-N-formylcolchicine, colchicine, cornigerine, 2-demethyldemecolcine, 3-demethyldemecolcine and demecolcine) were isolated from seven *Colchicum species*. However, although many alkaloids have been identified in *Colchicum*. The major alkaloid of *Colchicum* is colchicine. All parts of *Colchicum species* have been shown to contain colchicine, but seeds and corms contained more colchicine than other plant parts<sup>(61-64)</sup>.

***Conium maculatum***

*Conium maculatum* contained piperidine alkaloids (coniine, N-methyl-coniine, conhydrine, pseudoconhydrine, gamma-coniceine), which were formed by the cyclisation of an eight-carbon chain derived from four acetate units. Gamma-coniceine was the precursor of the other hemlock alkaloids. All vegetative organs, flowers and fruits contained alkaloids. The concentrations (both absolute and relative) of the different alkaloids depend on plant varieties, on ecological conditions and on the age of the plant. Conmaculatin (2-Pentylpiperidine), a novel volatile alkaloid related to coniine was identified from the *Conium maculatum* L. Chemical analysis showed that all tissues of *Conium maculatum* were very rich in alkaloids, fruits being the richest with up to 1% (w/w) alkaloid, but the amount and mutual ratio of the several different alkaloids depends on plant variety, ecological conditions and the stage of phenological development<sup>(65-69)</sup>.

### ***Convolvulus arvensis***

*Convolvulus arvensis* was found to contain the tropane alkaloids, tropine, pseudotropine, tropinone as well as cuscohygrine, meso-cuscohygrine and calystegines<sup>(70-73)</sup>.

### ***Crotalaria juncea***

Chodesmine alkaloids, 074% of toxic dehydropyrrolizidine alkaloids (DHPAs) (isohemijunceines 0.05%, trichodesmine 0.016%, and junceine 0.008%) were isolated from *Crotalaria juncea*<sup>(74)</sup>.

### ***Cupressus sempervirens***

*Cupressus sempervirens* contained 0.7% alkaloids<sup>(75-76)</sup>.

### ***Cynodon dactylon***

*Cynodon dactylon* contained alkaloids 0.1%<sup>(77-78)</sup>.

### ***Dactyloctenium aegyptium***

*Dactyloctenium aegyptium* leaf extract contained alkaloids  $0.540 \pm 0.083$  mg/g dry weight<sup>(79-80)</sup>.

## **Datura Species**

Tropane alkaloids are a group of more than 200 compounds best known for their occurrence in the family *Solanaceae* comprising over 100 genera and 3000 plant species. Tropane alkaloids characterized by a two-ringed structure with a pyrrolidine and a piperidine ring sharing a single nitrogen atom and two carbon atoms. The amino group, typical for all alkaloids, is in most cases methylated. The most important natural tropane alkaloids are (-)-hyoscyamine and (-)-scopolamine (also known as hyoscine). High concentrations of these alkaloids have been found particularly in *Datura* species. Hyoscine [(-)-Scopolamine] represented the main tropane alkaloid in *Datura fastuosa* (*Datura metel*). However, the plant contained alkaloids, hyosyamine, hyoscine and atropine. The total alkaloid content of the leaves was 0.426% which were mainly atropine. The seeds contained 0.426% and the flower contained 0.43% hyoscyamine. Scopolamine and atropine contents in the whole plant of the plant increased gradually with the progress of developmental growth, and were most pronounced when the plant was at the end of its reproductive stage. The highest percentage of scopolamine accumulation in the root was after 16 weeks. The root was the organ which often accumulated higher amounts of atropine. The aerial parts, if compared with the root of the plant, usually accumulated relatively higher amounts of scopolamine and relatively lower amounts of atropine<sup>(81-84)</sup>.

Production of tropane alkaloids in *Datura stramonium* plants was found to start from the end of the second week after seed germination. The rates of atropine and scopolamine production were similar (0.05%) at this stage. The quantity of alkaloids reached maximum at the end of the tenth week after seed germination, then gradually decreased as the plants entered the generative phase. Alkaloid content depended on the plant part and the stage of plant growth. Leaves and capsules showed the highest alkaloid content in the vegetative and generative phases, respectively. Generally the younger parts of plants contained more alkaloids than older ones. Alkaloid content decreased rapidly in leaves in the generative phase. Scopolamine was lowest (0.013%) in roots in the vegetative period, and then totally disappeared in the generative period. Atropine occurred in roots in both the vegetative (0.045%) and generative (0.056%) periods. Stems were rich in atropine (0.070%) but poor in scopolamine (0.023%) in both stages<sup>(84)</sup>.

The maximum contents of atropine in different parts of *Datura stramonium* were found in the stems leaves and seeds. The maximum contents of hyoscyamine and scopolamine in different parts of *Datura stramonium* were found in the stems and leaves of young plants, hyoscyamine being always the predominant component. In the Young plants, the maximum atropine level was recorded in the stems and medium leaves  $0.915 \pm 0.015$  and  $0.831 \pm 0.014$   $\mu\text{g}/\text{mg}$  respectively, while in adult plant, the maximum level of atropine was recorded in the flowers and small leaves  $0.270 \pm 0.026$  and  $0.165 \pm 0.006$   $\mu\text{g}/\text{mg}$  respectively. On the other hand, in the Young plants, the maximum scopolamine level was recorded in the stems and flowers  $0.129 \pm 0.014$  and  $0.106 \pm 0.031$   $\mu\text{g}/\text{mg}$  respectively, while in adult plant, the maximum level of scopolamine was recorded in the seeds and flowers  $0.089 \pm 0.010$  and  $0.066 \pm 0.004$   $\mu\text{g}/\text{mg}$  respectively<sup>(85-86)</sup>. However, sixty-four tropane alkaloids have been detected in *Datura stramonium* these included: Hygrine; 3 $\alpha$ ,6 $\alpha$ -Ditigloyloxy-7-hydroxytropine; 6-Hydroxyhyoscyamine; Pseudotropine; 3 $\alpha$ -Tigloyloxytropine; Hydroxy-6-tigloyloxytropine; Phenyl acetoxytropine; 3-Tigloyloxy-6-(2-methylbutyryloxy) tropine; Hyoscyamine; 3-Tigloyloxy-6-isovaleroyloxy-7-Hydroxytropine; Scopolamine; Tropinone; Scopine; 6-Hydroxyacetoxytropine; 3,6-Diacetoxytropine; 3-Tigloyloxy-6-acetoxytropine; 3-Tigloyloxy-2-methyl butyryl oxytropine; 3 $\alpha$ ,6 $\alpha$ -Ditiglotoxytropine; 3-Acetoxy-6-isobutyryloxytropine; 3-(2-Phenylpropionyloxy) tropine; Littorine; 6-Hydroxy apoatropine; 3 $\alpha$ ,6 $\alpha$ -Ditigloyloxy-7-hydroxytropine; 3-Tropoyloxy-6-acetoxytropine; 3,6-Dihydroxytropine; 3 $\alpha$ -Tigloyloxytropine; 3-Tigloyloxy-6-propionyloxy-7-hydroxytropine; 3 $\alpha$ -Apotro-poyloxytropine; Aposcopolamine; 3 $\alpha$ ,6 $\alpha$ -Ditigloyloxytropine; 3-(3'-Acetoxytropoyloxy) tropine; 3 $\alpha$ -Tigloyloxy-6-hydroxyt tropine; Tropine; 3-Acetoxytropine; 3-Hydroxy-6-acetoxytropine; 3-Hydroxy-6-methylbutyryloxytropine; 3-Tigloyloxy-6-isobutyryloxytropine;

Aponorscopolamine; 7-Hydroxyhyoscyamine; Meteloidine; 3 $\alpha$ ,6 $\alpha$ -Ditigloyloxytropine; 3-phenylacetoxy-6, 7-epoxynortropine; 7- hydroxyapoatropine and scopoline<sup>(84-84)</sup>.

#### ***Delphinium ajacis***

The total alkaloidal content of leaf, stem and root of the plant was 3.2, 1.9 and 4.2% w/w respectively, while seeds contained 1.01–1.06% alkaloids<sup>(89-91)</sup>. The alkaloids delsoline, acetyl delcosine, delcosine, gigactonine, 14-deacetylajadine, ajacine, ambiguine and ajadinine were isolated from the seeds of *Delphinium ajacis*<sup>(92-93)</sup>. Norditerpenoid alkaloids, ajadelphine, ajadelphinine, delcosine, delsoline, deltaline, gigactonine, 18-methoxygadesine and delphisine were isolated from the roots of *Delphinium ajacis*<sup>(94)</sup>.

#### ***Delphinium brunonianum***

*Delphinium brunonianum* contained norditerpenoid and diterpenoid alkaloid (anthriscifoldine)<sup>(95)</sup>. The norditerpenoid alkaloids which were divided into two main structural groups; the highly toxic MSAL-type and the less toxic MDL-type. Plants high in the MSAL-type alkaloids are thought to be the most toxic to cattle and the concentrations of these alkaloids have been used as a predictor of the plant toxicity<sup>(96-98)</sup>.

#### ***Ephedra species***

Ephedra species contain alkaloids ephedrine, pseudoephedrine, norephedrine, norpseudoephedrine, methylephedrine, and methylpseudoephedrine. Beside the E-type alkaloids, ephedroxane, and macrocyclic spermidines called ephedradine A-D, which isolated from some Eurasian Ephedra species. The total amount of alkaloids isolated from *Ephedra alata* aerial parts was 0.2-0.22%<sup>(99-101)</sup>.

#### ***Equisetum arvense***

Alkaloids such as nicotine, palustrine and palustrinine were isolated from the plant<sup>(102-103)</sup>.

#### ***Eryngium creticum***

The total alkaloids in the plant reached  $0.57 \pm 0.0058$  wt %<sup>(104-105)</sup>.

#### ***Eschscholzia californica***

Both aerial parts and roots contained alkaloids, the latter being richer than the former, it contained up to 1.6% alkaloids. The alkaloids most commonly reported in *Eschscholzia californica* included sanguinarine, dihydrosanguinarine chelirubine, macarpine, dihydromacarpine, californidine, chelerythrine, dihydrochelerythrine, chelilutine, dihydrochelilutine, sanguirubine, escholtzine, N-methylaurotetanine, caryachine, O-methylcaryachine, protopine, allocryptopine, reticuline, methyl laurotetanine, hunnemanine, norsanguinarine, pavine alkaloid (6S,12S-neocaryachine-7-O-methyl ether N-metho salt), 1-(3-hydroxy -4-methoxybenzyl) -2-methyl-6,7- methylenedioxy- 1,2,3,4-tetra hydro-isoquinoline, as well as the dihydro-intermediates. The amounts of alkaloids isolated from the aerial part of *Eschscholzia californica* were: protopine :  $0.514 \pm 0.038$ , californidine:  $12.5 \pm 1.8$ , allocryptopine:  $0.0120 \pm 0.0023$ , eschscholtzine:  $8.700 \pm 0.51$ , sanguinarine:  $0.0191 \pm 0.0050$ , chelerythrine:  $0.068 \pm 0.011$ , reticuline:  $1.095 \pm 0.16$ , n-methylaurotetanine:  $5.68 \pm 0.72$  and caryachine:  $0.410 \pm 0.065$  mg/ kg dry weight<sup>(106-114)</sup>.

#### ***Eupatorium cannabinum***

Pyrrolizidine alkaloids of the *Eupatorium cannabinum* were included echinatine isomers, lycopsamine and intermedine, and a number of their beta-acetyl, beta-angelyl/tiglyl and beta-(iso)valeryl esters. Pyrrolizidine alkaloids without a substituent at C-7 were tentatively identified as supinine and amabiline. In addition to a number of these alkaloids, some beta-(iso)butyryl, beta-angelyl/tiglyl, and beta-(iso)valeryl esters of supinine or amabiline were detected in subterranean parts of the plant. Pyrrolizidine alkaloids with a saturated necine base trachelanthamine isomers and some beta-anglyl/tiglyl esters which were detected in the root material only. A C-9- viridifloryl/trachelanthyl ester of a saturated amino-alcohol like turnefordicine and one of its beta-angelyl/tiglyl esters have also been found<sup>(115-118)</sup>.

#### ***Ficus carica***

*Ficus carica* contained crude alkaloid 9.6% /100g dry weight<sup>(119-120)</sup>.

#### ***Ficus semicordata***

Alkaloid content of the plant leaves was ranging from  $18.91 \pm 0.16$  to  $45.68 \pm 0.55$  mg/gm. The methanolic extract showed more alkaloid content  $45.68 \pm 0.55$  mg/gm than other extracts<sup>(121)</sup>.

#### ***Fritillaria imperialis***

The bulb was poisonous raw, it contained low concentrations of a toxic alkaloid(12). As well as imperialine, two other alkaloids of molecular formula  $C_{27}H_{41}NO_2$  and  $C_{27}H_{45}NO_3$  were obtained from bulbs of *Fritillaria imperialis* var. *rubra maxima*<sup>(122)</sup>. A new class of C-nor-d-homo steroidal alkaloids (impranane), impranine and dihydroimpranine, a new pyridyl-pregnane-type steroidal alkaloid, fetisinine and the base korsevine were isolated from the bulbs of *Fritillaria imperialis*<sup>(123)</sup>.

Ebeinone isolated from the bulbs of *Fritillaria imperialis* exhibited anticholinergic activity and completely blocked the responses to acetylcholine<sup>(124)</sup>. Ebeinone at concentration of (1 $\mu$ g/ml) exhibited anticholinergic activity as manifested by blocking of acetylcholine response in isolated guinea pig ileum and atria<sup>(125)</sup>.

The steroidal bases (impericine, forticine, delavine, persicanidine A, and imperialine) isolated from the ethanol extract of the air-fried bulbs of *Fritillaria imperialis* possessed inhibitory effect on compounds causing anti-acetylcholinesterase and anti-butrylcholinesterase activity. In order to check the structure-activity relationship and prepare more potent derivatives of imperialine with anticholinergic activity, imperialinol, 3 beta-acetoxyimperialine, 3 beta-propionoxyimperialine, and 3 beta-butyroxyimperialine were prepared. 3 betapropionoxyimperialine, and 3 beta-butyroxyimperialine displayed better anticholinergic activity against muscarinic receptors of the heart and brain than imperialine. The decrease in activity in imperialinol showed the importance of the 6-keto functionality in imparting the anticholinergic activity<sup>(126)</sup>.

The ability of the alkaloid, ebeinone, isolated from *Fritillaria imperialis*, for binding with muscarinic M2 and M3 acetylcholine receptors was investigated. In functional studies with guinea-pig left atrium, ebeinone was found to be 10-fold more active as an antagonist of responses to carbachol (CCh) than in either guinea-pig ileum or trachea. The estimated dissociation constants (KB values) in the three tissues were 77.3, 931.1 and 547.0 nM, respectively. Inhibition binding studies in rat atria with the non-selective antagonist [3H]N-methylscopolamine ([3H]NMS) showed that ebeinone had a KI value of 80.9 nM. Comparison of ebeinone with pancuronium showed that both compounds able to retard the dissociation rate of [3H]NMS in atria, indicating an allosteric mode of interaction at the M2 receptor<sup>(127-128)</sup>.

Imperialine (cervane alkaloid), was assessed at M1, M2 and M3 receptors in functional assays and at M1, M2, M3 and putative M4 sites in binding studies. In functional assay, imperialine appeared as a selective surmountable antagonist at M2 receptors in guinea-pig isolated atria and uterus ( $-\log KB = 7.7$  and  $7.4$ , respectively), in comparison to M1 receptors in canine isolated saphenous vein ( $-\log KB = 6.9$ ) or M3 receptors in a range of guinea-pig isolated smooth muscles including ileum, trachea, fundus, seminal vesicle or oesophagus ( $-\log KB = 6.6-6.8$ ). In rat aorta, the  $-\log KB$  value at the M3 receptor ( $5.9$ ) was slightly, but significantly, lower. In competition radioligand binding studies, imperialine was also selective toward M2 sites in rat myocardium ( $-\log Ki = 7.2$ ) with respect to M1 and M3 sites (rat cerebral cortex, rat submaxillary gland;  $-\log Ki = 6.1$  and  $5.7$ , respectively). However, it did not significantly discriminate between rat cardiac M2 sites and putative M4 sites in rabbit lung ( $-\log Ki = 6.9$ )<sup>(129)</sup>.

### ***Fumaria officinalis***

The plant contained alkaloids isoquinoline-type. Protopines including protopine (fumarine) as the major alkaloid and cryptopine, protoberberines including aurotensine, stylophine, sinactine and N-methylsinactine. Two new isoquinoline alkaloids, fumaranine and fumarostrejdine, along with 18 known alkaloids were isolated from aerial parts of *Fumaria officinalis*<sup>(130)</sup>.

Total alkaloids was 540 mg/ 100g dry weight of the aerial parts of plants, alkaloids isolated from *Fumaria officinalis* were included : protopine (protopine: 42 mg/100g dry weight, cryptopine: 11 mg/100g dry weight), tetrahydroprotoberine (sinactine: 6 mg/100g dry weight), (adlumine: 2 mg/100g dry weight) and spirobenzyl isoquinoline (parfumine: 2 mg/100g dry weight, fumariline: 3 mg/100g dry weight, fumarophycine: 3 mg/100g dry weight, fumaritine: < 2 mg/100g dry weight)<sup>(131)</sup>. However, the weight of protopine alone could be reach 230 mg/ 100 g of *Fumaria officinalis* dry weight<sup>(132-133)</sup>.

### ***Fumaria parviflora***

The plant contained wide range of alkaloids, the methanolic extraction of 10.5 kg of the plant yielded 0.39% of total alkaloids, the following alkaloids have been identified parfumine, norjusiphine, Nmethyladlumine, d-fumaricine, adlumiceine, adlumidiceine, (+)- adlumidine, (+)-adlumine, (-)-adlumine, (+)-bicuculline, ( $\pm$ )-bicuculline, bicucullinine (narceimine), (-)-cheilanthifoline, coclaurine, coptisine, (-)-corlumine, cryptopine, dehydrocheilanthifoline, dihydrodrohmariline, dihydrosanguinarine, fumaramidine, fumaramine, fumaridine, fumariflorine ethyl ester, fumarilicine, (+)-fumariline, fumaritine, (+)- $\alpha$ -hydrastine, (+)-isoboldine, izimirine, lahoramine, lahorine, 8- methoxydihydrodrohmariline, methylhydrastinium, N-methyladlumine, N-methylhydrasteine, N-methylhydrastine, N-methylhydrohydrasteine, narlumidine, norjuziphine, noroxy- hydrastinine, 8-oxocoptisine, oxysanguinarine, ( + )- pahidine, (+)-parfumine, (+)-parviflorine, protopine, thequaternarysalt of protopine, sanguinarine, (-)-scoulerine, ( - )-stylophine, and ( $\pm$ )-stylophine<sup>(134-139)</sup>.

Phthalide isoquinoline alkaloid [( $-$ )-corlumine] and rhoeadines-like alkaloid [rhoeagenine] were isolated from *Fumaria parviflora*<sup>(138, 140)</sup>.

The Total alkaloids mg/ 100g dry weight of the aerial parts of plants was 521, the quantities of different alkaloids isolated from *Fumaria parviflora* were: protopine (protopine: 57 mg/100g dry weight, cryptopine: 5 mg/100g dry weight), tetrahydroprotoberine (sinactine: 2 mg/100g dry weight), (adlumine: 3 mg/100g dry weight) spirobenzyl isoquinoline (parfumine: 14 mg/100g dry weight, fumariline: 10 mg/100g dry weight, parfumidine: 7 mg/100g dry weight) and benzophenanthridine (dihydrosanguinarine: 2 mg/100g dry weight)<sup>(141-142)</sup>.

### **Galium aparine**

*Galium aparine* seeds contained 2.76±0.03% alkaloids<sup>(143-144)</sup>.

### **Glaucium corniculatum**

Many alkaloids were isolated from *Glaucium corniculatum*, the phytochemical analysis revealed that *Glaucium corniculatum* of Czechoslovakian origin contained protopine, allocryptopine, corydine, isocorydine, chelirubine, chelidonine, chelery- thrine, sanguinarine, coptisine, berberine, stylopine methohydroxide and canadine methohydroxide. *Glaucium comiculafum* Rud. subspecies refractum from Iran contained dicentrine (0.20%), glaucine (0.18%), dicentrinone (0.17%), dehydrodicentrine (0.05%), (±)-chelidonine (0.07%), α-allocryptopine (0.02%), protopine (0.02%), and N-methylaurotetanine (0.01%). However, in Egyptian species, the three major alkaloids were, protopine, corydine and isocorydine. The four minor ones were condine N-oxide, isocorydine N-oxide, N-methylcorydine and dihydroprotopine. Protopine, allocryptopine, corydine, isocorydine, chelidonine, berberine were isolated from *Glaucium corniculatum* of Iraqi origin. *Glaucium corniculatum* of Bulgaria origin contained protopines: (protopine, allocryptopine, cryptopine); aporphines: (corydine, isocorydine, norglucine, glaucine, N-methylaurotetanine, isoboldine, corunnine); benzophenantridines: (Chelitytryne, Sanguinarine, Chelidonine); protoberberines: (canadine, berberine) and spirobenzylisoquinolines: (corydaine). While, *Glaucium corniculatum* of Algerian origin contained protopines: (protopine, allocryptopine); aporphines: (corydine, glaucine); benzophenantridines: (Chelidonine); protoberberines: (canadine, berberine); benzylisoquinolines: (berbitine) and spirobenzylisoquinolines: (corydaine). Glaucine, chelidonine, protopine, allocryptopine and coptizine alkaloids were isolated from *Glaucium corniculatum* from Iasi<sup>(145-150)</sup>.

### **Glossostemon bruguieri**

*Glossostemon bruguieri* root contained alkaloids (5.8 ± 0.43% of dry raw weight)<sup>(151-153)</sup>.

### **Gossypium hirsutum**

*Gossypium hirsutum* contained alkaloids 12.20± 0.28%<sup>(154-155)</sup>.

### **Haplophyllum Species**

*Haplophyllum buxbaumii* contained pyranoquinoline type alkaloid<sup>(156)</sup>. Alkaloid (+)-tuberine, haplotubinone and haplotubine were also isolated from *Haplophyllum tuberculatum*<sup>(156-159)</sup>.

Quinoline alkaloids and lignan lactones, included: quinoldione, 3-(1',1'- dimethylallyl) -3-(3'',3''-dimethylallyl) -1,2,3,4-tetrahydro-2,4- quinoldione, 4-(3',3'-dimethylallyloxy) -3-(3'',3''-dimethylallyl)-2(1H)- quinolone, Polygamain, kusunokinin and 1-methyl-2-n-nonyl-4(1H)- quinolone were isolated from *Haplophyllum tuberculatum*. Haplotubinone and haplotubine alkaloids, were also isolated from the aerial parts of *Haplophyllum tuberculatum*<sup>(159-161)</sup>. The aerial parts of *Haplophyllum tuberculatum* collected in Sudan yielded the furoquinoline alkaloid skimmianine<sup>(162)</sup>. Chemical analysis of *Haplophyllum tuberculatum* naturally growing in Saudi Arabia revealed the presence of amide alkaloids, tuberine, tubacetine, tubasenicine and 7-Hydroxy-4-Methoxy-8-prenylfuro[2,3- b]quinolone<sup>(163-165)</sup>.

### **Hedera helix**

Alkaloid (emetin) was isolated from *Hedera helix*<sup>(166-167)</sup>.

### **Helianthus annuus**

Quantitative phytochemical analysis of ethanolic leaf extract of *Helianthus annuus* showed that it contained alkaloid 1.23%<sup>(168-169)</sup>.

### **Heliotropium Species**

Four pyrrolizidine alkaloids (heleurine, heliotrine, supinine, and europine) were isolated from *Heliotropium bacciferum*. The pyrrolizidine alkaloids isolated from the aerial plant parts of *Heliotropium europaeum* were included: supinine-type 2.3 ± 0.4 µg/g dry weight (included: supinine, supinine-N-oxide, heleurine, heleurine-N-oxide); Heliotrine-type 80.0 ± 7.9 µg/g dry weight (included: heliotrine, europine, heliotrine-N-oxide, europine-N-oxide, rinderine, 5'-hydroxyrinderine, 3'-acetyl rinderine, rinderine-N-oxide, 5'-hydroxyrinderine-N-oxide, 3'-acetyl rinderine-N-oxide, 5'-acetyeuropine, 5'-acetyeuropine-N-oxide, echinatine-N-oxide); and Lasiocarpine-type 28.7 ± 2.2 µg/g dry weight (included: 7-angeloylheliotrine-N-oxide, lasiocarpine, lasiocarpine-N-oxide, iso-lasiocarpine, iso-lasiocarpine-N-oxide, 5'-acetyl lasiocarpine, 5'-acetyl lasiocarpine-N-oxide, isoacetyl lasiocarpine, iso-acetyl lasiocarpine-N-oxide, heliosupine, 3'-acetylheliosupine, heliosupine-N-oxide and 3'-acetylheliosupine-N-oxide). The total pyrrolizidine alkaloid and tertiary base content of the seeds of *Heliotropium europaeum* were found to be 0.28 % and 0.02 % respectively. Higher percentage of alkaloids were present as N-oxides (92.86 % of the alkaloids). Alkaloids found in the tertiary base fraction and total alkaloid fraction were identified as europine 14.27%, heliotrine 2.44%, supinine 9.09%, heleurine 2.65%, lasiocarpine 8.69% and 7-angelylheliotrine 2.86%. *Heliotropium europaeum* population Garmsar was shown to contain three major alkaloids: heliotrine N-oxide 0.08%, lasiocarpine 0.09% and lasiocarpine N-oxide 0.05%; and four minor alkaloids: heliotrine 0.02%, europine 0.02%,

acetylasiocarpine 0.03% and a novel alkaloid acetylasiocarpine N-oxide 0.05%. Alkaloid, heliotridine esterified on the methylol hydroxyl with lasiocarpic acid (2,3-dihydroxy-4-methoxy-2-methylpentane-3-carboxylic acid), supinine (supinidine esterified with trachelanthic acid), supinidine esterified with heliotric acid (3-hydroxy-4-methoxy-2-methylpentane-3-carboxylic acid), N-oxides of heliotrine and lasiocarpine were isolated from *Heliotropium europaeum*<sup>(170-176)</sup>.

#### ***Hibiscus rosa-sinensis***

The flower extract of *Hibiscus rosa-sinensis* (Red) contained 0.51±0.16 % alkaloids. While , the flower extract of *Hibiscus rosa-sinensis* (White) contained 0.50±0.18 % alkaloids and the flower extract of *Hibiscus rosa-sinensis* (Yellow) contained 0.48±0.16 % alkaloids<sup>(177)</sup>.

#### ***Hibiscus sabdariffa***

*Hibiscus sabdariffa* contained alkaloids 2.14%<sup>(178-180)</sup>.

#### **Hyoscyamus species**

All *Hyoscyamus* species are rich sources of tropane alkaloids, mainly hyoscyamine and scopolamine<sup>(181)</sup>. The total alkaloid content was 1%. Eighteen alkaloidal compounds were detected, in hairy root cultures of *Hyoscyamus albus* obtained by an infection of plants with *Agrobacterium rhizogenes*, six of the compounds were in trace amounts. However, some of these alkaloids were hygrine, tropinone, tropine, pseudotropine, 3 $\alpha$ -tigloyloxytropane, 3  $\beta$ -acetoxytropane, cuscohygrine, apoatropine, hyoscyamine, littorine, scopolamine and 6  $\beta$ -hydroxyhyoscyamine<sup>(182-184)</sup>.

Thirty-four alkaloids were identified in the roots of *Hyoscyamus albus*, 23 in the stems, 24 in the leaves, 24 for the flowers and 21 in the seeds. The identified alkaloids included: hygrine, cyclopropine, tropinone, tropine, pseudotropine, scopoline, scopine, 2,5-(2-oxopropyl)-hygrine (2,5-diacetyl-N-methylpyrrolidine), 3-(hydroxyacetoxyl) tropane, N-methylpyrrolidinyl- hygrine A, N-methylpyrrolidinyl-hygrine B, 3 $\alpha$ - tigloyloxytropane, cuscohygrine, 3 $\beta$ -tigloyloxytropane, 6,7-dehydro-3- phenyl acetoxytropane, 3-phenylacetoxyl tropane, 3-(2'-phenylpropionyl)oxy tropane (dihydroapoatropine), 6,7-dehydro-3-apotropyloxytropane, apohyoscyamine, phygrine, 3-phenylacetoxyl-6,7- epoxytropane, 6,7-dehydrohyoscyamine, 3-(3'-methoxy tropoyloxy) tropane, 3-phenylacetoxyl-6-hydroxytropane, aponor scopolamine, aposcopolamine, N-methylpyrrolidinyl-cuscohygrine A, N-methyl pyrrolidinyl-cuscohygrine B, hyoscyamine (atropine), 6-hydroxyapo hyoscyamine, scopolamine, 4'-hydroxylittorine, 7-hydroxyhyoscyamine and 6-hydroxy hyoscyamine. Hyoscyamine (atropine) was the major

isolated alkaloid, represented 63.8, 77.8, 70.2, 66.3 and 80.4% of the alkaloids isolated from the roots, stems, leaves, flowers and seeds respectively, followed by scopolamine which represented 4.2, 9.1, 16.6, 16.5 and 6.4% of the alkaloids isolated from the same parts respectively<sup>(185)</sup>.

The main tropane alkaloid of the *H. reticulatus* plant was hyoscyamine in the range from 0.033 to 0.056% dry weight, followed by scopolamine from 0.011 to 0.015% dry weight<sup>(186)</sup>. The quantitative analysis of *Hyoscyamus reticulatus* from Iran showed that it contained 0.031% hyoscyamine and 0.025% scopolamine. The total alkaloid content of the leaves of *H. reticulatus* from Turkey was found to be in the range of 0.011-0.027%<sup>(184, 187)</sup>. The maximum hyoscyamine and scopolamine concentrations were found in the leaf, and minimum concentration in the stem. Total alkaloids in the leaf, stem and capsule were 0.7126 , 0.2099 and 0.3686 mg/g respectively, and the total alkaloids in the leaf and root of cultured plant were 5.0844 and 0.8556 mg/g respectively. In leaf, stem and capsule of collected plants, hyoscyamine concentrations were 0.3515, 0.0788 and 0.3192 mg/g, and scopolamine concentrations were 0.3611, 0.1311 and 0.0494 mg/g respectively. However, in leaf and root of cultured plant, hyoscyamine concentrations were 2.3377 and 0.1683 mg/g, and scopolamine concentrations were 2.7467 and 0.6873 mg/g respectively<sup>(188)</sup>.

From the *in vitro* hairy root cultures of *Hyoscyamus reticulatus*, 10 tropane alkaloids were identified (hydrin, tropin,  $\alpha$ -acetyltropin, 11- acetyltropin, cuscohygrin, apoatropin, littorin, hyoscyamine, scopolamine, 6- $\beta$ -hydroxyhyos cyamine), and 4 from normal plant roots (apoatropin, littorin, hyoscyamine, scopolamine). However, the content of hyoscyamine and scopolamine in the leaves and roots of normal plants was maximal in and before flowering stage, and the maximal foliar scopolamine content was before flowering. In the roots the changes in hyoscyamine and scopoamine production were not marked as in the leaves. The changes in the content of both main alkaloids in *Hyoscyamus reticulatus* were greater in the leaves than in the roots during different growth periods<sup>(189)</sup>.

#### ***Lagerstroemia indica***

Biphenyl and biphenyl ether quinolizidine N-oxide alkaloids were also isolated from the plant. Decamine, decinine, decodine, dihydroverticillatine, lagerstroemine and lagerine alkaloids were isolated from *Lagerstroemia indica*<sup>(190-193)</sup>.

***Lantana camara***

*Lantana camara* leaves contained 9.76±0.02 mg/g alkaloids<sup>(194-195)</sup>.

***Lawsonia inermis***

Two alkaloids, harmine and harmaline were also isolated from the ethanol extract of *Lawsonia inermis* leaves<sup>(196-197)</sup>.

***Leontice leontopetalum***

*Leontice leontopetalum* contained alkaloids 7.4 ±0.32 mg/g to 27.12 ±1.18 mg/g dry weight (leontidine, leontine, leontamine, lupanine, 13 $\alpha$ -hydroxy lupanine,  $\alpha$ - isolupanin, 3 $\alpha$ - hydroxylupanin, leontiformidine, d-leontiformine, pachycarpine, oblongine, petaline, (+) O-methyldihydro secoquettamine and (+) dihydrosecoquettamine) and up to 30% starch. The plant also contained flavones and saponins with a hemolytic index of 1:240 in the aboveground portion of the plant<sup>(198-201)</sup>. The alkaloid pattern of *Leontice leontopetalum* was characterized by quinolizidine alkaloids of the lupanine-type with lupanine as the main compound. In *Leontice leontopetalum* L. subsp. *ewersmannii* 15 quinolizidine alkaloids were detected, in contrast to *Leontice leontopetalum*, *L. ewersmannii* accumulated quinolizidine alkaloids of the matrine-type and the  $\alpha$ -pyridone-type was the major compounds<sup>(198-200)</sup>.

***Lippia nodiflora***

*Lippia nodiflora* contained 0.589% alkaloids<sup>(201-203)</sup>.

***Lithospermum officinale***

Two pyrrolizidine alkaloids, O-7- 3-hydroxy-3-methylbutanoyl-O-9-(-)-hydroxy viridifloryl retronecine and its acetyl derivative were isolated from *Lithospermum officinale*<sup>(204-205)</sup>.

***Lolium temulentum***

Analysis of the overground part of *Lolium temulentum* led to the isolation of two main alkaloids: loline and perloline. Loline and two minor alkaloids (2-hydroxy -7- methoxy-1-methylamino and the corresponding 1-aminopyrrolizidine) were strictly localized in the caryopses, whereas perloline was found mainly in the stems<sup>(206)</sup>.

***Luffa acutangula***

The fruit of *Luffa acutangula* contained 0.19 mg/kg alkaloids<sup>(207-208)</sup>.

***Lycium barbarum***

Two steroidal alkaloid glycosides were isolated from the seeds of *Lycium barbarum*<sup>(209)</sup>.

***Lythrum salicaria***

Different parts of *Lythrum salicaria* contained: alkaloids (piperidine and quinolizidine derivatives: lythranine, lythranidine, lythra mine, lythrancineI–VII, lythrancepineI–III)<sup>(210-212)</sup>.

***Mangifera indica***

The mango stem bark and leaves contained 9.66 ± 0.20 and 0.84 ± 0.11 mg/ 100g alkaloids<sup>(213)</sup>.

***Marrubium vulgare***

Alkaloids (emetine and cephaeline) were isolated from *Marrubium vulgare*<sup>(214)</sup>.

***Medicago sativa***

*Medicago sativa* contained include alkaloids (stachydrine, homostachydrine)<sup>(215)</sup>.

***Mirabilis Jalapa***

*Mirabilis Jalapa* leaves contained 0.034mg/kg alkaloids<sup>(216-217)</sup>.

***Oxalis corniculata***

Quantitative phytochemical analysis of *Oxalis corniculata* showed that total alkaloids of the plant was (0.86 mg/kg)<sup>(218-219)</sup>.

***Narcissus tazetta***

Many alkaloids: pretazettine, tazettine, homolycorine, haemanthamine, ismine, narcisine, narciclasine, lycorine, pseudolycorine, pseudolycorine N-oxide, galanthamine, nor-galanthamine, 11- hydroxygalanthine, buphanisine, haemantamine, 3-epihydroxy bulbispermine O- methylmaritidine, 9-O-Demethylhomolycorine, 3-epi hydroxybulbispermine and N-methyl-8,9-methylenedioxyphenanthridinium were isolated from *Narcissus tazetta*<sup>(220-224)</sup>.

## II. CONCLUSION

Medical use of alkaloid-containing plants has a long history and many alkaloids are still used in medicine nowadays. The current manuscript discussed the plant contents of alkaloids for medical, pharmaceutical, synthetic, and many other useful properties.

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