Medicinal plants affected male and female fertility (part 1)- A review

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Abstract:- The previous studies revealed that many medicinal plants were affected male and female fertility. These plants included: Achillea santolina, Ailanthus altissima, Alhagi maurorum, Allium cepa, Althaea rosea, Ammarnia baccifera, Anethum graveolens, Anthemis nobelis, Arctium lappa, Asplenium trichomanes, Avena sativa, Bacopa monniera, Bryophyllum calycinum, Caesalpinia crista, Calendula officinalis, Calotropis procera, Capsella bursa-pastoris, Carthamus tinctorius, Carum carvi, Chenopodium album, Cicer arietinum, Cistanche tubulosa, Citrullus colocynthis, Citrus species, Coriandrum sativum, Cressa cretica, Crocus sativus, Crotalaria juncea, Cuminum cyminum, Cynodon dactylon, Cyperus rotundus, Dactylotcenium aegyptium, Dalbergia sissoo, Datura fastuolsa, Datura metel, Daucus carota, Dodonaea viscoso, Euphorbia hirta, Ficus carica, Phoenix dactyliphera. This review will highlight the medicinal plants which affected male and female fertility.

Keywords: Medicinal plants, herbs, fertility, reproductive systems, male, female

I. INTRODUCTION

Plants are a valuable source of a wide range of secondary metabolites, which are used as pharmaceuticals, agrochemicals, flavours, fragrances, colours, biopesticides and food additives. As a result of accumulated experience from the past generations, today, all the world’s cultures have an extensive knowledge of herbal medicine. Two thirds of the new chemicals identified yearly were extracted from higher plants. 75% of the world’s population used plants for therapy and prevention. In the US, where chemical synthesis dominated the pharmaceutical industry, 25% of the pharmaceuticals were based on plant-derived chemicals [1]. The previous studies showed that a wide range of synthetic drugs and medicinal plants exerted many effects on reproductive systems function [2-26]. These medicinal plants affected male and female fertility were included: Achillea santolina [27], Ailanthus altissima [28], Alhagi maurorum [29], Allium cepa [30], Althaea rosea [31], Ammarnia baccifera [32], Anethum graveolens [33], Anthemis nobelis [34], Arctium lappa [35], Asplenium trichomanes [36], Avena sativa [37], Bacopa monniera [38], Bryophyllum calycinum [39], Caesalpinia crista [40], Calendula officinalis [41], Calotropis procera [42], Capsella bursa-pastoris [43], Carthamus tinctorius [44], Carum carvi [45], Chenopodium album [46], Cicer arietinum [47], Cistanche tubulosa [36], Citrullus colocynthis [48], Citrus species [49], Coriandrum sativum [50], Cressa cretica [51], Crocus sativus [52], Crotalaria juncea [53], Cuminum cyminum [54], Cynodon dactylon [55], Cyperus rotundus [56], Dactylotcenium aegyptium [56], Dalbergia sissoo [56], Datura fastuolsa [56], Datura metel [56], Daucus carota [56], Dodonaea viscoso [56], Euphorbia hirta [57], Ficus carica [57], Phoenix dactyliphera [58-59]. This review was designed to highlight the medicinal plants which affected male and female fertility.

I- MEDICINAL PLANTS AFFECTED MALE REPRODUCTIVE PERFORMANCE

<table>
<thead>
<tr>
<th>Plant</th>
<th>Effects</th>
<th>Ref.</th>
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<tbody>
<tr>
<td><strong>Achillea santolina</strong></td>
<td>hydroalcoholic extract (300 mg/kg/day ip. for 20 days) caused histological alterations in the seminiferous tubules included disorganized germ epithelium, exfoliation of immature germ cells, germ cell necrosis and increased number of metaphases in germinal epithelium of seminiferous tubules in mice.</td>
<td>60</td>
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<tr>
<td><strong>Ammannia baccifera</strong></td>
<td>ethanol extract of A. baccifera whole plant induced antifertility effects in rat males. It was significantly reduced the weight of the testis, epididymis, sperm density and motility, content of fructose in the seminal vesicles, Δ5-3β -hydroxy steroid dehydrogenase (Δ5-3β-HSD) and glucose-6-phosphate dehydrogenase (G-6-PD)</td>
<td>61</td>
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<tr>
<td><strong>Arctium lappa</strong></td>
<td>roots extract at 600 and 1,200 mg/kg body weight significantly increased the frequencies of mount, intromission, and ejaculation frequency (p &lt; 0.05). Administration of the extract also reduced the post-ejaculatory interval.</td>
<td>62 - 64</td>
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<tr>
<td><strong>Bacopa monniera</strong></td>
<td>caused reversible suppression of spermatogenesis and fertility. The treatment caused reduction in motility and viability of the sperms and reduced the number of spermatozoa in cauda epididymis and testis, and caused alterations in the somniferous tubules in mice.</td>
<td>65</td>
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<tr>
<td><strong>Caesalpinia crista</strong></td>
<td>caused morphological changes in the sperm of albino rats including disturbance in the plasma membrane and acrosomal membrane. Considerable changes in the shape and size of the sperm head were observed, with the middle region of the sperm head being slightly constricted dorso-ventrally. Most sperm appeared morphologically abnormal in the head region showing the distortion at the anterior region and bulging of the acrosomal membrane when compared with the control.</td>
<td>66</td>
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<tr>
<td><strong>Carthamus tinctorius</strong></td>
<td>Induced formation of multinucleated giant cells in the germinal epithelium. It also caused a significant decrease in seminiferous tubule diameter, seminiferous epithelium height and maturation arrest (p&lt;0.001).</td>
<td>67</td>
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<tr>
<td><strong>Chenopodium album</strong></td>
<td>ethanolic extract at doses of 100, 250 and 500mg/kg orally, in male albino mice induced significant increase in the mount frequency, intromission frequency, intromission latency as well as aggregate of penile reflexes and significant reduction in the post ejaculatory interval. Moreover 500 mg/kg, orally, was found to be the most effective dose.</td>
<td>68</td>
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<td>The ethanolic extract of seeds at a concentration of 200 mg/kg bw resulted in pronounced anabolic effect in treated male rats as evidenced by an increased body weight as well as the weight of reproductive organs. Sexual behavior and performance were also markedly improved as reflected in reduction of mount, intromission and post ejaculatory latency. Furthermore, the extract also enhance sperm count. seed extract induced sperm death, the effect which is due to oxidative damage of cellular macromolecules by generation of ROS.</td>
<td>69 - 70</td>
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<td>Aqueous decoction of <em>Chenopodium album</em> seeds (CAD) was assessed for its sperm-immobilizing and contraceptive efficacy in laboratory mammals. The minimum effective concentration of CAD that induced instantaneous immobilization of rat spermatozoa in vitro was 2 mg/ml. The mechanism of CAD action involved disintegration of sperm plasma membrane and dissolution of acrosomal cap causing sperm death.</td>
<td>71</td>
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<tr>
<td><strong>Cicer arietinum</strong></td>
<td>Oral administration of methanolic extract at 200 and 400 mg/kg body weight was significantly increased the mount frequency, intromission frequency, ejaculation frequency and ejaculation latency (P &lt; 0.05) in rats. It also significantly (p&lt;0.05) increased the serum cholesterol and testosterone levels.</td>
<td>72</td>
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<tr>
<td>Medicinal Plant</td>
<td>Effect</td>
<td>Reference</td>
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<tr>
<td><strong>Cistanche tubulosa</strong></td>
<td>Administration of ethanol extract (0.4 and 0.8 g/kg) increased sperm count (2.3 and 2.7 folds) and sperm motility (1.3 and 1.4 folds) and decreased the abnormal sperm (0.76 and 0.6 folds) in rats respectively. The serum level of progesterone and testosterone in rats was also increased by CTE administration (p&lt;0.05). Results of immunohistochemistry and western blot analysis confirmed that the expression of CYP11A1, CYP17A1, and CYP3A4 was enhanced by CTE (p&lt;0.05). The weights of seminal vesicle and prostate gland of castrated young rats were significantly increased by administration of alcohol soluble extract from the decoction of <em>Cistanche tubulosa</em>.</td>
<td>73</td>
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<td><strong>Citrullus colocynthis</strong></td>
<td>50% ethanol extract caused significant reduction of cauda epididymis sperm motility and density, number of pups, fertility, and circulatory levels of testosterone were observed in male rats. The weights of testes, epididymis, seminal vesicle, and prostate were also significantly decreased. The concentration of testicular cholesterol was significantly elevated, while protein, sialic acid, acid and alkaline phosphatase concentrations were decreased. The histoarchitecture of the testes showed degenerative changes in the seminiferous epithelium, arrest of spermatogenesis at the secondary spermatocyte stage, cytolysis, and the lumen filled with eosinophilic material.</td>
<td>74</td>
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<td><strong>Citrus species</strong></td>
<td>Lime juice destroys sperm cells, fifty percent of <em>Citrus aurantifolia</em> juice wiped out 2000 of sperm cells in 30 seconds. 60 days treatment with <em>Citrus limonum</em> seeds significantly decreased the sperm count. Size and weight of testis and epididymis were reduced indicating atrophic changes in testis and epididymis. It caused drastic effect on sperm motility and morphology which decreased fertility. Sperm counts returned to normal after 90 days lime juice caused reduction in the number of fetus of treated pregnant rats when compared to the control. There was a significant reduction in the crown-rump length, weight and umbilical cord length of the fetus when compared with the control. Accordingly, lime juice showed abortificient effect but no obvious teratogenic effect was observe.</td>
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<td><strong>Cressa cretica</strong></td>
<td>Methanolic extract of <em>Cressa cretica</em> 100 mg/kg for 60 days led to a significant decrease in the weight of testis, epididymis, seminal vesicle, and ventral prostate. <em>Cressa cretica</em> reduced the fertility of male rats by 100%. There was a marked reduction in the number of primary spermatocytes, secondary spermatocyte, and spermatids. Sertoli cell counts were significantly decreased. Leydig cell nuclear area and the number of mature Leydig cells were also significantly decreased. Various fractions of the methanol extract for 60 days decreased the weight of testes and accessory sex organs significantly (P≤0.001). Sperm counts of testes and cauda epididymis as well as cauda epididymal sperm motility was also declined significantly (P≤0.001), serum testosterone production was reduced in treated male rats. The fertility was decreased by 90%-100% in different fractions. The seminiferous tubular diameter and Leydig cell nuclear area were reduced significantly. The population of spermatogenic cells</td>
<td>76</td>
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(spermatogonia, preleptotene, pachytene, secondary spermatocytes and round spermatids) were also reduced significantly.

<table>
<thead>
<tr>
<th>Medicinal plants</th>
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<tr>
<td><strong>Crocus sativus</strong></td>
<td>The aqueous extract (80, 160 and 320 mg/kg bw), crocin (100, 200 and 400 mg/kg bw), increased mounting frequency, intromission frequency and erection frequency behaviors and reduced ejaculation latency, intromission latency and mount latency parameters. The prepared saffron gel significantly improved erectile dysfunction in diabetic patients (P &lt; .001)</td>
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<td><strong>Crotalaria juncea</strong></td>
<td>Plant extracts were tested in male mice and rats, they decreased the weights of testis and accessory reproductive organs. The diameters of the testis and seminiferous tubules were decreased. Spermatogonia, spermatocytes and spermatids in the testis and the sperm count in cauda epididymis were also decreased. Ethanol extract appeared to be the most potent antispermatogenic extract. When the ethanol extract was tested in immature male mice, it exerted antiandrogenic effect as the weights of accessory organs were reduced</td>
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<tr>
<td><strong>Cuminum cyminum</strong></td>
<td>Oral dose of Cuminum cyminum isolated fractions (CcFr) 50 mg/rat/day for 60 days caused marked abnormalities in spermatogenesis with decreased counts (P ≤ 0.001) in round spermatids, preleptotene spermatocytes and secondary spermatocytes. Cross sectional surface area of Sertoli cells as well as number of mature Leydig cell were decreased significantly (p≤0.001). Testicular as well as accessory sex organ biochemical parameters were significantly changed (p≤0.001). Sperm motility, density and morphology were resulted in 100% negative fertility. Testosterone levels were declined significantly.</td>
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<td><strong>Dactyloctenium aegyptium</strong></td>
<td>The extract was administered orally by gavage in the dose of 500 and 800 mg/kg bw per day as a single dose in male rats for 28 days. It increased mounting frequency and the mating performance of the rats highly significantly (p&lt;0.01). The extract also influenced the behaviour of treated animals in comparison to non-treated rats in a remarkable manner, making them more attracted to females</td>
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After 60 days oral administration of Cressa constituents, results showed 100% antifertility activity in male rats with the reduction in testosterone levels and spermatogenic elements.
vesicle and prostate, with a significant reduction of total sperm count and increase in motility, abnormality of sperm in caput and caudal were also recorded. Histologically, the treated groups showed dose related reduction in the diameter of seminiferous tubules, with reduced layering, less spermatозoa, hyper-cellularity of leydig cells with the presence of large multinucleated Cells. The administration of ethanolic extract of D. aegyptium to males also showed dose dependent decrease in number of pregnant females and number of fetuses.

**Dalbergia sissoo**

Ethanolic extract of stem bark caused dose-dependent and time-dependent adverse effect on sperm motility and sperm viability. Ethanol extract at a concentration of 20 mg/ml caused complete immobilization within 3 minutes. Ethanol extract at a dose of 200 mg/kg in mice resulted in a significant decrease (p<0.001) in weight of the testis and epididymis. A significant decrease (p<0.01) in sperm motility and sperm count in the epididymis were also observed with pathohistological changes.

**Datura fastuolsa**

Alcoholic extract of (2, 4 and 6 mg/kg, for 7 weeks) in male rats induced significant decrease in concentrations of sperm and normal sperm in all the concentrations in. They also significantly decreased serum levels of testosterone, libido of treated males showed no change, but their fertility was markedly suppressed.

**Daucus carota**

Administration of carrot seed extract (CSE) caused a significant increase in cauda epididymis sperm reserves compared with the control (28.2 ± 1.8 ×10^6 vs. 45.1 ± 2.0, ×10^6). The extract also protect testis from the gentamicin-induced necrosis. The CSE administration caused about 3.5-times increase in the LH levels even in spite of receiving 5 mg/kg/day gentamicin with no significant effect on FSH levels. The testosterone concentrations in the group received 400 mg/kg CSE were 30% and 83% higher than its levels in the control and the gentamicin treated group, respectively.

**Dodonaea viscosa**

Leaf extracts showed antifertility activity in male rats. It decreased sperm count and reproductive organ weights with the appearance of necrotic changes in the seminiferous tubules of testis. Total protein and glycogen levels were reduced in treated rats.

**Euphorbia hirta**

Aqueous extracts (400 mg/kg orally) in old mature male rats caused varying degrees of testicular degeneration and reduction in the mean seminiferous tubular diameter. *E. hirta* exerted potentially induced deleterious effects on the tested and accessory organs of rats.
An aqueous ethanol extract of the dried fruits of *Ficus carica* was screened for in vivo aphrodisiac activity. Results reveal that on the 1st day of treatment all the treated groups showed increase copulatory sexual behavior and orientational activity in all the experimental animals. The prolonged treatments for all the treated groups were highly effective for increase the sexual libido, as compared to the solvent control.

The effect of *Ficus carica* leaf extracts 200 mg/kg, in sperm parameters and testis was studied in mice intoxicated with formaldehyde. The results showed that formaldehyde significantly decreased gonadosomatic index and increased percentage of immotile sperm. Disorganized and vacuolated seminiferous epithelium, spermatogenic arrest, and lumen filled with immature germ cells were also observed in the testes of mice intoxicated with formaldehyde. However, leaf extracts improved sperm count, nonprogressive motility of spermatozoa, and gonadosomatic index in formaldehyde-treated testes.

Pollen of Date palm (500 mg iq) and a combination of zinc sulphate & pollen of Date palm (500 mg iq) in infertile men significantly increased serum LH, FSH, & testosterone levels. It was also, increased significantly sperm count and motility. Sexual desire was also significantly increased. Wives of treated men got pregnancy during the treatment period.

### II. III-MEDICINAL PLANTS AFFECTED FEMALE REPRODUCTIVE PERFORMANCE

<table>
<thead>
<tr>
<th>Plant</th>
<th>effects</th>
<th>Ref</th>
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<tbody>
<tr>
<td><em>Ailanthus altissima</em></td>
<td>was recognized to have anti-progestogenic activities. It inhibited the progesterone activity in a dose-response manner</td>
<td>97</td>
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<tr>
<td><em>Alhagi maurorum</em></td>
<td>ethanolic extract of powdered roots in doses of 5 mg/ml bathing fluid completely suppressed histamine induced uterine contractions</td>
<td>98</td>
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<tr>
<td><em>Allium cepa</em></td>
<td>female rats treated with ethanolic extract showed significant inhibition of number of implant sites at a dose of 300 mg/kg. It enhanced uterine contraction in rats equivalent to 0.003 IU of oxytocin.</td>
<td>99</td>
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<tr>
<td><em>Althaea rosea</em></td>
<td>infusion and methanolic extract influence hormonal activity and affected the morphology of the rat female sexual organs. It exerted estrogenic activity, the <em>in vivo</em> test proved that <em>p</em>-hydroxy benzoic acid isolated from the plant was estrogenic.</td>
<td>100</td>
</tr>
<tr>
<td><em>Ammannia baccifera</em></td>
<td>ethanol extract at the doses of 100, 200 and 400 mg/kg body weight (ip) arrested the normal estrus cycle at dioestrus phase and significantly decreased weight of ovaries. The cholesterol and ascorbic acid content in ovaries were significantly elevated in treated mice. The extract also significantly inhibited the activity of Δ5-3β-hydroxy steroid dehydrogenase and Glucose-6-phosphate dehydrogenase, the two key enzymes involved in ovarian steroidogenesis</td>
<td>101</td>
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<tr>
<td><em>Anthemis nobelis</em></td>
<td>aqueous-alcoholic extract was studied in polycystic ovary syndrome induced in rats by a single dose of estradiol valerate. Histological investigations revealed that the animal administered with dose of 50 mg/day showed small cysts and less inflammation, with decreasing of serum estrogen hormone</td>
<td>102</td>
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<tr>
<td>Medicinal plants</td>
<td>Effects and Actions</td>
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<tr>
<td><strong>Anethum graveolens</strong></td>
<td>0.045 g/kg and 0.45 g/kg of aqueous extract and 0.5 g/kg and 5 g/kg of ethanol extract for 10 days caused significant increase in duration of the estrous cycle and diestrus phase. Smooth endoplasmic reticulum (SER), rough endoplasmic reticulum (RER) and mitochondria were increased in granulosa lutein cells. Dill seed possessed contractive effects on myometer, enhanced releasing of oxytocin which is an effective hormone in uterus contractions. A dose of 6-7 gm of dill seed extract after delivery decreases postpartum hemorrhage due to its contractive characteristic. Limonene and anethole showed contractive effect on uterine myometrium. Smooth endoplasmic reticulum (SER), rough endoplasmic reticulum (RER) and mitochondria were increased in granulosa lutein cells. Contractive effects in the treated women (one tablespoon of whole dill seed seeped in a half or whole cup boiling water for 3-4 min before going to the hospital at the beginning of uterus contractions) was significantly more than the control group. The ratio of contraction’s fall time to its rise time in the treated group was shorter than the control group. The study showed that dill seed shortens duration of the first stage of labor.</td>
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<td><strong>Arachis hypogaea</strong></td>
<td>Introduction of refined peanut oil to form 10% of the food ration of immature mice increases uterine weight. Plant showed high levels of phytoestrogens including isoflavones (formononetin and biochanin A, 729 ug/g dry weight. These compounds structurally or functionally mimic mammalian estrogens.</td>
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<td><strong>Arctium lappa</strong></td>
<td>Induced uterine stimulant activity</td>
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<td><strong>Asplenium trichomanes</strong></td>
<td>In vitro estrogenic activity to activate ERalpha and ERbeta. MCF7/ERE1uc cell line which expresses endogenous ERalpha and SK-NBE cells transiently transfected with the estrogen receptors (ER alpha and ER beta) were used to test the estrogenic activity assays. Leaves infusion and methanolic extract were active in MCF7 model; selectivity for the ERbeta receptor was observed in the SK-NBE test.</td>
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<td><strong>Avena sativa</strong></td>
<td>Oat straw stimulated the release of luteinizing hormone from the adenohypophysis of rats. It contained oestrone which been shown to induce ovulation.</td>
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<td><strong>Bryophyllum calycinum</strong></td>
<td>Exerted relaxant effect in vitro on the contractility of human myometrium on oxytocin-stimulated contraction at a minimum concentration almost 100-fold lower than in the case of spontaneous contraction. Thirty-two patients divided into two groups. 15 patients received Bryophyllum and 17 received the placebo. The time of delivery did not differ between the groups. In both groups A transition to the intensive care unit was slightly higher in the placebo group (13) compared to the Bryophyllum group (11).</td>
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<td><strong>Caesalpinia crista</strong></td>
<td>It caused antifertility effect in mice and rats. This effect could be attributed to its contents of gossypol and cyclopropane fatty acids, which recognized as an antifertility compounds. Alcoholic seed extract caused histological follicular degeneration in ovary, vacuolation and mild disorganization of uterus in rats. It also caused significant decrease in duration of estrous cycle and mean ovarian weight.</td>
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<tr>
<th>Medicinal Plant</th>
<th>Effect</th>
<th>Reference</th>
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<tr>
<td>Calendula officinalis</td>
<td>extracted exerted estrogenic activity in ovariectomized animals</td>
<td>123-125</td>
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<tr>
<td>Calotropis procera</td>
<td>The effects of ethanolic and aqueous extracts were found to interrupt the normal oestrous cycle in 60% and 80% of female rats respectively. The extracts had no oestrogenic activity when tested in immature female bilaterally ovariectomized rats.</td>
<td>126</td>
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<td>A strong antiimplantation (inhibition 100%) and uterotrophic activity was possessed by ethanolic extract at the dose level of 250 mg/kg (1/4 of LD50).</td>
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<td>Aqueous extracts induced significant sustained increases in human myometrial smooth muscle cell contractility, with varying efficiencies, depending upon time of exposure and dose.</td>
<td>128</td>
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<tr>
<td>Carum carvi</td>
<td>aqueous and ethanolic extracts showed significant antifertility activity in female rats. FSH and LH levels were significantly decreased, and estrogen was found to be increased. The estrus phase was blocked by treatment with aqueous and ethanolic extract. They also increase the weight of ovary, uterus and body weights.</td>
<td>129</td>
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<td>Oil was effective in inhibiting tonic and phasic rhythmic contractions of isolated uterine preparations</td>
<td>130</td>
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<tr>
<td>Capsella bursa-pastoris</td>
<td>Impeded ovulation and produced temporary infertility in males and females</td>
<td>131</td>
</tr>
<tr>
<td>Carthamus tinctorius</td>
<td>In studying the teratogenic effect, in higher doses (1.6 and 2 mg/kg/day) the embryos were absorbed, whereas with lower dose (1.2 mg/kg/day) it caused changes in external, internal and longitudinal diameters, open neuropore, changes in cellular orientation and cellular degeneration were observed.</td>
<td>132</td>
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<td>Tracheloside isolated from the plant, significantly decreased the activity of alkaline phosphatase (AP), an estrogen-inducible marker enzyme, with an IC50 value of 0.31 microg/ml, a level of inhibition comparable to that of tamoxifen (IC50=0.43 microg/ml).</td>
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<td>Exerted stimulating action on the uterus of mouse in vitro. The stimulating action of Carthamus tinctorius has been found related to the stimulating effects on H1-receptor and alpha-adrenergic receptor of uterus. Intraperitoneal administration of a hot aqueous extract flowers increased uterine contractions in pregnant female rats</td>
<td>134</td>
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<tr>
<td>Chenopodium</td>
<td>Fertilization of oocytes and establishment of implantation were</td>
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<tbody>
<tr>
<td>Album</td>
<td>Prevented in the uterine horn that was administered with CAD. In rabbit, intravaginal application of CAD significantly blocked the establishment of pregnancy. Accordingly, CAD possesses appreciable spermicidal potential, which may be explored as an effector constituent of vaginal contraceptive.</td>
<td>19</td>
</tr>
<tr>
<td>Cicer arietinum</td>
<td>The aqueous extract at a dose of 400mg/kg was found to be most effective abortifacient. Similarly, it was also found to increase the reproductive organ weight and possess estrogenic activity when tested in immature ovariectomised female albino rats. Isoflavones extracted from chickpea sprouts (ICS) stimulated estrogen responsive element (ERE)-promoter activity in cells, and concurrent treatment with the nonselective estrogen receptor antagonist ICI 182,780 abolished the estrogenic activity induced by ICS. Treatments of rats with isoflavones extracted from chickpea sprouts (ICS) (50 or 100 mg/kg/day) produced significant estrogenic effects on the uteruses, including the increases in uterine weight, epithelial height and gland number, as well as in the expression of the cell proliferation marker PCNA. The treatments changed the secretory profile of ovarian hormones and pituitary gonadotropins: (serum E2 level was significantly increased, while serum LH and FSH levels were decreased).</td>
<td>137, 138, 139</td>
</tr>
<tr>
<td>Citrullus colocynthis</td>
<td><em>Citrullus colocynthis</em> 400 mg/kg in female rats for 4 weeks did not have much effect on fertility. Significant decrease in the relative ovarian weights and embryo weights in female rats exposed to <em>Citrullus colocynthis</em> were observed. Exposure to <em>Citrullus colocynthis</em> for a 12 weeks resulted in a reduction in the percentage of pregnancies and in the number of implantation sites, decreased ovarian weights and decreased viable fetus's number.</td>
<td>140</td>
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<tr>
<td>Citrus species</td>
<td>Petroleum ether extract of seeds of <em>Citrus medica</em> in female rats showed reduced ovarian weight, benzene extract treated rats showed increased ovarian weight, ethanol extract treated rats showed non-significant change in the weight of ovary. Histological changes of the ovary indicated increases in the number of atretic follicles but decreases in the number of healthy developing follicles, Graafian follicles and corpora lutea. The total cholesterol, activity of acid and alkaline phosphatase and ascorbic acid content of the ovary were increased. Petroleum ether extract of <em>Citrus medica</em> seeds exhibited estrogenic effects, which included increase in uterine weight and vaginal epithelial cell cornification in female rats. The micrometric measurements of the uterus and its components were increased and glands showed high secretory activity. When the extract was tested in 30-day-old immature rats, they exhibited opening of vagina on the fifth day and cornification of vaginal epithelial cells, which was about 10 days earlier compared to controls. Petroleum ether extract of <em>Citrus medica</em> leaves proved to retain high estrogenic activity in immature female rats.</td>
<td>141, 142, 143</td>
</tr>
<tr>
<td>Coriandrum sativum</td>
<td>Aqueous extract of fresh seeds produced a dose-dependent significant anti-implantation effect, but did not produce complete infertility in female rats. Treatment of animals</td>
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A review

During day-8 to day-12 and day-12 to day-20 of the pregnancy did not produce any significant abortifacient activity. There was no significant change in the weight and length of the foetuses delivered by rats treated with the extract and no abnormalities were seen in the organs of the offsprings. The extracts produced a significant decrease in serum progesterone levels on day-5 of pregnancy which may be responsible for its anti-implantation effect.

### Crocus sativus

Saffron aqueous extract (SAE), was evaluated in *in vitro* maturation (IVM) of immature mouse oocytes. The maturation rate was significantly higher in all groups treated with different concentrations of SAE compared with the control group (p<0.05). However, the lower concentrations of SAE (10 and 5 µg/ml in maturation medium) increased the fertilization rate of oocytes and *in vitro* developmental competence when compared with the control group (p<0.05).

The effects of different concentrations of saffron (*Crocus sativus*) aqueous extract (SAE) and its ingredient, crocin, were evaluated on the improvement of *in vitro* maturation (IVM) and subsequent *in vitro* fertilization (IVF) and embryo development of mouse oocytes. SAE was added at dosages of 5, 10, and 40 µg/ml and crocin 50, 100, and 400 µg/ml. Both SAE and crocin improved the rate of IVM, IVF, and *in vitro* culture. Addition of 40 µg/ml SAE to maturation medium significantly increased the rate of IVM, IVF, and *in vitro* culture (p < 0.05). Furthermore 100 µg/ml crocin significantly increased the IVM rate (p < 0.05).

A double-blind and placebo-controlled trial was designed to investigate the effect of saffron (stigma of *Crocus sativus*) on the symptoms of premenstrual syndrome. The trial showed that saffron was effective in relieving symptoms of PMS. A significant difference was observed in efficacy of saffron in the total premenstrual daily symptoms and Hamilton depression rating scale.

### Crotalaria Juncea

Petroleum ether, benzene and alcohol extracts of seeds of *Crotalaria juncea* administered orally at the dose level of 25mg/100g bw to adult female mice for 30 days, resulted in irregular estrous cycle with prolonged estrus and metaestrus and reduced diestrous and proestrus during the experimental period. Histological studies of the ovary indicated increases in the number of atretic follicles but decreases in the number of developing follicles, Graafian follicles and corpora lutea.

Ethanol extract of *Crotalaria juncea* seeds which showed antiovulatory activity in female albino rats. Two fractions decreased number of healthy follicles (Class I – ClassVI) and corpora lutea and increased number of regressing follicles (Stage IA, Stage IB, Stage IIA, Stage IIB).

Alcohol extract possessed antiimplantation and pregnancy interruption activities. These adverse effects on fertility were reversible upon withdrawal of the extract treatments. The alcohol extract was found to possess estrogenic activity.
| **Cynodon dactylon** | aqueous extract of entire plant of *Cynodon dactylon* for thirty days in female rats significant increased (p<0.001) the serum estradiol concentration whereas, follicle stimulating and luteinizing hormones were significantly (p<0.001) reduced. Furthermore, a significant increase (p<0.001) in the weight of the uterus and significant decrease in the weight of the ovaries (p<0.001) was observed in the treated group, the estrous cycle was found to be irregular and disturbed | 151-152 |
| **Cyperus rotundus** | the essential oil of the rhizome of *Cyperus rotundus* (EOC ) and its fractions F2-F6 showed significant anti-dysmenorrhea. | 153 |
| **Datura metel** | 2% acetone seed extracts for 15 days in female mice caused 100% anti-implantation activity followed by 1% and 0.5% seed extracts which caused 40% and 80% anti implantation activity respectively. | 154 |
| **Daucus carota** | petroleum ether extract and fraction 5 (fatty acids) of carrot seeds arrested the normal estrus cycle of adult mouse and reduced the weight of ovaries significantly. The cholesterol and ascorbic acid content in ovaries were significantly elevated by the extract and fraction 5 of carrot seeds. The significant inhibition of delta 5,3-beta-hydroxy steroid dehydrogenase and glucose-6-phosphate dehydrogenase, the two key enzymes involved in ovarian steroidogenesis, were also recorded in mouse ovaries after 15 days of treatment The petroleum ether, alcoholic, and aqueous extracts of *Daucus carota* were evaluated for their possible antiovulatory activity in rabbits with copper-induced ovulation. All extracts inhibited ovulation in 40%, or less, of the animals. | 155 |
| **Dodonea viscose** | methanolic extract of the leaves reduced significantly (p<0.01) the number of liters when administered through oral route in female rats. It also produced anti-fertility effect in a dose dependent manner and the contraceptive effect was manifested for a definite period of time. Furthermore, the extract significantly showed anti-implantation and early abortifacient activity | 156 |

### III. CONCLUSION:

The paper reviewed the effects of the medicinal plants on the functions of reproductive systems in males and females, to be utilize in medical applications as a result of effectiveness and safety.
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