

CHATURBEEJA IN PRIMARY DYSMENORRHOEA (KASHTARTAVA): AN OBSERVATIONAL STUDY

OBSERVATIONAL STUDY Dhiman Kamini^{1*}, Chaudhary Kiran²

¹Reader, Deptt of SRPT, IPGTRA, GAU, Jamnagar (Gujarat) India

²Lecturer, Deptt of SRPT, Govt. Ayu. College, Riva (M.P.) India

*Email: kd44ayu@yahoo.co.in

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ABSTRACT

In folk medicines, Chaturbeeja (combination of seeds of four plants i.e Trigonella Foenum-graecum, Lepidium sativum, Nigella sativa, Trachyspermum ammi in equal quantity) has been traditionally used for variety of applications including treatment of Dysmenorrhoea, the most common gynaecological symptom reported by women. To promote the proper use of such medicines and to determine their potential as sources for new drugs, it is essential to study medicinal plants, which have folklore reputation in a more intensified way. A Single blind, prospective observational clinical study was conducted to evaluate the efficacy of Chaturbeeja powder in Primary Dysmenorrhoea (Kashtartava). 25 patients were administered Chaturbeeja powder in a single dose of 3g with hot water at night, 7 days before starting of menstruation till 3rd day of the menstruation cycle. After assessing the results it was observed that12 patients were markedly improved, 8 moderately improved and 5 were improved.

Keywords: Dysmenorrhoea, Kashtartava, Chaturbeeja, Trigonella Foenum-graecum, Lepidium sativum, Nigella sativa, Trachyspermum ammi, Folk medicine.

INTRODUCTION:

India is well known historically as a land of spices and aromatic plants and continues to be one of the leading producers of spices and medicinal plants in the world. The beneficial health effects of many plants, used for centuries as seasoning agents in food and beverages, have been claimed for preventing various ailments. To promote the proper use of such medicines and to determine their potential as sources for new drugs, it is essential to study medicinal plants, which have folklore reputation in a more intensified way. Chaturbeeja churna/powder has long been a part of the folk medicine from many parts of the world for the complaint of Dysmenorrhoea. Dysmenorrhoea is of two types out of which Primary dysmenorrhea is characterized by a cramps in/spasmodic pain in supra pubic pain that begins somewhere between several hours before and a few hours after the onset of the menstrual bleeding. Symptoms peak with maximum blood flow and usually last less than one day, but the pain may persist up to 2 to 3 days². The pain is characteristically colicky and located in the midline of the lower abdomen but may also be described as dull and may extend to lower quadrants, the lumbar area, and the thighs. Frequently associated symptoms include diarrhoea, nausea and vomiting, fatigue, light-headedness, headache, dizziness and, rarely, syncope and fever³⁻⁵. Dysmenorrhea is the most common gynaecological symptom reported by women. Ninety percent of women presenting for primary care suffer from some menstrual pain⁶. Population surveys suggest that, although prevalence rates vary considerably by geographical location, complaints of dysmenorrhea are widespread in diverse populations⁷⁻¹¹. Furthermore, one third to one half of these women report moderate or severe symptoms. Symptoms are frequently associated with time lost from school, work, or other activities¹².

Kashtartava is the term which is being used for the condition where in a woman may suffer with pain during menstruation and it covers all the problems & ailments that are mentioned in various artavavyapat/Yonivyapada & other diseases related with female reproductive system in classical literature of Ayurveda. By analysis one can state this is a result of

vitiation of Apaana vata (pelvic physiology regulator), Apanavatamargavarodha (disturbed pelvic physiology), and artavadushti (menstrual dysfunction) & dhatukshava (tissue loss). Kashtartava can be compared with the Dysmenorrhea in modern medicine. In Ayurvedic clasics no separate management is mentioned independently Kashtartava(Dysmenorrhoea). Ayurveda advocates many a medicament for the management of this problem as per the disease condition. Several research works had been reported in this regard also. In this study an attempt has been made to evaluate the efficacy of a purely compound herbal formulation i.e Chaturbeeja Churna (combination of seeds of four plants i.e Methi- Trigonella Foenum-graecum, Chandrashur -Lepidium sativum , Kalajaji- Nigella sativa and Yavanika- Trachyspermum ammi in equal quantity),) highlighted in *Ayurvedic* text *for* its efficacy in *Kashtartava*/Primary Dysmenorrhoea¹³⁻¹⁹.

MATERIALS AND METHODS:-

Collection and preparation of Drug:-

Dried and cleaned raw seeds of *Methi (Trigonella Foenum-graecum), Chandrashur (Lepidium sativum), Kalajaji (Nigella sativa), Yavanika (Trachyspermum ammi)* were purchased from local market after being authenticated by the Dravyaguna experts of the college. The seeds were powdered with a mechanical grinder to pass through a 0.8-mm mesh sieve and stored in an airtight glass container.

Patients: 25 (twenty five) Patients, attending the OPD of *Prasooti tantra* and *Stree roga* at Rajiv Gandhi Govt. Ayurvedic College Hospital, Paprola Dist. Kangra (HP), with characteristic features of *Kashtartava* (Dysmenorrhoea), were selected and registered irrespective of their caste, creed, religion, income, occupation for the present study.

Study Design: An observational study.

PROTOCOL DURING TRIAL:-

- i. Fulfilment of inclusion criteria.
- ii. Consent of patient after making her aware of the merits/demerits of the trial.
- iii. Registration of the patient.
- iv. Investigations done before inclusion into the trial.

- v. Follow up of the patient every month for assessment and clinical evaluation.
- vi. Data so available and deducted clinically was statistically analysed.

CRITERIA OF INCLUSION:-

- Patients coming with chief complaint of Dysmenorrhoea with scanty or average bleeding during periods.
- Age group between 12 40 years.
- Patients suffering for more than 6 menstrual cycles.

CRITERIA OF EXCLUSION:-

- Patients not willing for trial,
- Patients having congestive dysmenorrhoea
- Patients below 12 years and above 40 years,
- Patients with chronic general illness,
- Patients with intrauterine contraceptive devices.
- Patients having problem of Menorrhagia and any anatomic or uterine pathology - fibroid, adenomyosis were excluded from the study.

INVESTIGATIONS:-Routine investigations of blood and urine were carried out to rule out associated systemic pathology. Sonography (U.S.G.) was done for uterine and adenexal study, if needed, to rule out any pathology or lesion. **DRUG SCHEDULE:** Chaturbeeja Churna, 3g at bed time, orally, with Luke warm water for 10 days (starting 7 days before commencement of menstruation cycle till 3rd day of the of the bleeding phase) with Luke warm water in a dose of 3g twice a day (12 hourly) for two consecutive cycles.

DURATION OF TRIAL: The total duration of treatment for the subjects was 2 (two) months.

FOLLOW UP STUDY: Follow up was conducted after one month during trial and then after the completion of trial.

CRITERIA FOR ASSESSMENT: - The effect of treatment (results) was assessed regarding the clinical signs and symptoms (on the basis of VAS and grading, scoring system). Overall improvement was observed and recorded as Before Treatment (BT) and After Treatment (AT).

CLINICAL ASSESSMENT: The criteria adopted for intensity of pain was VAS (Visual Analogue Scale) which is a measurement instrument that tries to measure a characteristic or attitude that is believed to range across a continuum of values and which cannot be easily and directly measured. Operationally a VAS is usually a horizontal line, 100 mm (10 cm) in length, anchored by word descriptor at each end, as illustrated in figure-1.

The patient marks on the line the point that they feel represent their perception of their current state. The VAS score is determined by measuring in millimetres from the left hand end of the line to the point that the patient marks.

Other signs and symptoms were assessed by adopting suitable scoring methods. The details are illustrated in Table

Pain as bad

Overall score of each symptom was recorded as follows:-

Absence of symptom Presence in mild degree Presence in moderate degree Presence in severe degree

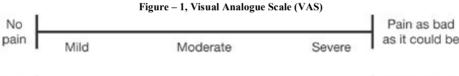




Table 1: Grading of signs and symptoms

GRADING	0	1	2	3
Intensity	No pain	Mild	Moderate	Severe
Duration	No pain	Up to 24hrs	24to <48hrs	48-<72hrs
Nausea	Absent	1-3times/day	4-5times/day	>5times/day
Vomiting	Absent	Occasionally	1-2times/day	>2times/day
Breast Tenderness	Absent	Mild	Moderate	Severe
Fever	Absent	Mild	Moderate	Severe
Headache	Absent	Mild	Moderate	Severe
Giddiness	Absent	Occasionally	1-2times/day	>2times/day
Diarrhoea	Absent	Occasionally	1-2times/day	>2times/day
Anorexia	Absent	Mild	Moderate	Severe
Nervousness	Absent	Mild	Moderate	Severe
Irritability	Absent	Mild	Moderate	
Constipation	Absent	Mild	Moderate	Severe
Weakness	Absent	Mild	Moderate	Severe
Bloating	Absent	Mild	Moderate	Severe

OVERALL RESULTS:-

Markedly improved ≥ 75%

Moderately improved 51%-75%, Improved 25%-50%,

<25% Unimproved

Statistical analysis: The obtained data on the basis of observations were subjected to statistical analysis in terms of mean, standard deviation, standard error and unpaired 't' test were conceded at the level of p<0.001 as highly significant, p<0.05 or p<0.01 as significant and p<0.10 or p>0.01 as insignificant to carry out the results.

RESULTS:

EFFECT OF THERAPY - The efficacy of Chaturbeeja powder orally in twenty five patients was noted and results derived after statistical analysis were as per the Table No. –

Table 2: Effect of chaturbeeja churna on signs and symptoms

Symptoms	Mean score		Relief		Paired -t	Paired –t test		
	B.T.	A.T.	Diff	% age	S.D.±	S.E. <u>±</u>	T	P
Intensity of pain	2.76	0.96	1.8	65.22	0.82	0.163	11.02	< 0.001
Duration of pain	1.75	0.76	0.99	56.57	0.71	0.141	7.07	< 0.001
Nausea	2.06	0.63	1.44	69.70	0.81	0.203	7.06	< 0.001
Vomiting	1.5	0.13	1.38	91.67	0.74	0.263	5.23	=0.001
Breast Tenderness	1.83	0.5	1.33	72.67	0.52	0.211	6.33	=0.001
Fever	1.14	0.43	0.71	62.48	0.49	0.184	3.87	< 0.01
Headache	1.5	0.5	1.0	66.67	0.47	0.149	6.71	< 0.001
Giddiness	1.78	0.71	1.07	60.02	0.83	0.221	4.84	< 0.001
Diarrhoea	1.56	0.66	0.89	57.24	0.33	0.111	8.00	< 0.001
Constipation	1.71	0.57	1.14	66.67	0.69	0.261	4.38	< 0.01
Anorexia	2.4	0.65	1.75	72.92	0.78	0.176	9.95	< 0.001
Nervousness	1.83	0.78	1.06	57.57	0.64	0.151	7.01	< 0.001
Irritability	2.0	0.91	1.09	54.55	0.92	0.196	5.55	< 0.001

OVERALL RESULTS: - 12 patients were markedly improved, 8 moderately improved and 5 were improved. **DISCUSSION**

This study was aimed to assess the efficacy of Chaturbeeja powder in the patients of Dysmenorrhoea. All the 25 registered Patients got significant improvement in almost all the symptoms.(table- 2) Duration and intensity of pain, tenderness in breast, headache reduced significantly. According to Ayurveda Vāta Prakopa is the main factor for Kashtartava (Dysmenorrhoea), the Vatashamaka (Pacifies Vata), Mridu Shodhana (do Purification softly), Sthapana (Relieve Pain), Shoola hara, actions of Chaturbbeeja due to their Snigdha Guna 50% and Ushna Virya (100%) with Vata-Kaphahara Dosha-karma (100%) may help to reduce the pain of the patients. Vitiation of Vata gets pacified by Sneha Guna and ushna virya as these are vata dosha suppressant due to having opposite qualities of vata. The individual ingredient of Chaturbeeja churna i.e Trigonella Foenum-graceum, Trachyspermum ammi, Lepidium sativum is reported for its antispasmodic, analgesic and estrogenic properties and this spasmolytic activity mediated through calcium channel blockade may relieves the pain by direct action on the myometrium. It has proven significant anti-oxidant activity, which by free-radical scavenging, enhances the immunity and general strength of the body. It increases the pain threshold and facilitates better pain tolerance capacity.

In the present study, significant improvement in the symptoms like nausea, vomiting, anorexia, diarrhoea, constipation was observed in all the patients.(table-2). In classical text of Ayurveda seeds of methi, chandrsura, kalajajee, and yavani are highlighted for their Tikta Rasa (75%) deepana, properties (Table-3) and indicated for Ajirna, agnimandya, conditions. Further the seeds of L sativum, T. foenum-graceum, T ammi, are reported for their antidiarrheal, germicide, antispasmodic, and antifungal agent. These drugs as shown in table 3 have Vata-shamaka, Deepana, Shoolahara, Jwarahara, Garbhashaya-shodhaka properties. Chaturbeeja Churna has pacified the vitiated Vata Dosha mainly due to Ushna Virya. Further, Laghu Guna (100%), Ruksha Guna (50%), pacified the slight Kapha vitiation. Hence, the properties of Chaturbeeja Churna can be made out as -Laghu Guna Katu Rasa - Katu Vipaka -Ushna Virya and Vata-Kaphahara. The drug mainly works with Ushna Virya as it is the most important property which determines the action of the drug. Individually, the drugs have the properties which help to cure dysmenorrhoea. The prepared Churna has bitter (Tikta) taste, thus having Mukhashodhaka and Agnivardhaka properties. So, it increases appetite, digestion and reduces nausea and vomiting. 20-20

Table 3: Pharmacological properties of chaturbeeja churna

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Dravya	Guna	Rasa	Vipaka	Virya	Dosha	Classical	Classical uses	Chemical	Pharmacological	
					karma	properties		composition	actions	
Methika	Laghu	Katu	Katu	Ushna	V. K.↓	Appetizer,	Anorexia, fever,	Steroidal	Anti -diabetic	
	Snigdha					Carminative,	diarrhea,	saponins		
	_					Emmenogogue.	Vomiting,	_		
						Astringent,	bloating.			
						Diuretic.	Menstrual			
							function.			
Chandrashura	Laghu,	Katu,	Katu	Ushna	$V. K \downarrow$.	Appetizer,	Low backache, in	Glycosides	Oestrogenic action,	
	Snigdha	Tikta				Diuretic and	rejuvenation.	Lepidimide	Anti-oxidant,	
						Emmenogogue.			Anti-spasmodic,	
									Diuretic.	
Kalajaji	Laghu,	Katu,	Katu	Ushna	$V.K\downarrow$.	Rajo-	Flatulence,	Alkaloids	Oestrogenic action,	
	Ruksha	Tikta			$P.\uparrow$	rodhanashaka,	Dysmenorrhoea.	Nigellimine	Anti-histaminic,	
						Diuretic, Ecbolic.			Anti-bacterial,	
									Spasmolytic.	
Yavani	Laghu,	Katu,	Katu	Ushna	V. K. ↓	Analgesic,	Flatulence,	Thymol	Inhibits platelet	
		Tikta			<i>P</i> .↑	Laxative.	abdominal pain.		aggregation, Muscle	
									relaxant, Anti -	
									oxidant.	

Thymol, the major phenolic compound present in Ajowan (*Trachyspermum ammi*), has been reported to be a germicide, antispasmodic, and antifungal agent. Numerous studies have

been carried out to reveal the therapeutic potential of fenugreek in various pathological conditions of gastric disorders 27-32 Diosgenin extracted from the seeds of

Foenum-graceum is used as a good antispasmodic, that it can be used for cramps, coughs and for muscular spasms. ^{33,34} *Trigonella Foenum-graceum* contains phyto-estrogen (which is a term applied to non-steroidal plant materials displaying estrogenic activity), It also possess sedative, anxiolytic, activity³⁵, ³⁶. *Lepidium* anxiolytic, myo-relaxant and analgesic sativum seed extract possesses antidiarrheal and spasmolytic activities.(due to mediated possibly through dual blockade of muscarinic receptors and Ca⁺⁺ channels). Trigonella foenum-graecum (TFG) is assumed to have a stimulating effect on digestive process. T. ammi has been used in cooking and as medicine, primarily to control indigestion and flatulence. It is prescribed for colic, diarrhoea, antibacterial and other bowel disorders.. Extracts prepared in different solvents exhibited variable activity against E. coli, P. aeruginosa, S. typhi and S. aureus suggesting their centuries old usage in the treatment of gastrointestinal disorders. This historical use of seeds to cure various gastrointestinal disorders has also been scientifically proved in another study carried out by Kaur and Arora (2009) antispasmodic and broncho-dilating actions in vitro. The studies on calcium channel blockade that has been found to mediate the spasmolytic effects of plant materials and considered that this mechanism contributed to their observed result and supported the traditional use of T. ammi in hyperactive disease states of the gut such as colic and diarrhoea as well as in hypertension³⁷⁻⁴⁷

In the present study relief was noted in the complaint of fever (table-2). The influence of the volatile oil of *Nigela sativa* on reducing the body temperature of mice has been reported. To followed by a detailed review with elucidation of the mechanism of action. In the present study relief was noted in the complaint of anxiety (table-2). *Nigella sativa* has been reported for its anti stress activity. Above all, N. sativa seed is a promising source for active ingredients which act through suppression of the inflammatory mediators prostaglandins and leukotriens.

The drug contains various minerals (Iron, Calcium, Phosphorus etc.) and vitamins (A, B, C) which are also necessary to maintain good health and proper functioning of the body systems. Thus, all these contribute to better health and improved psychology of the patients, allowing them better pain tolerance and even healthier stress-free life

SUMMARY & CONCLUSION:

Recent researches have also rationalized the uses of these drugs and further help us to scientifically prove the mode of action of the drug (Chaturbeeja Churna) in dysmenorrhoea. The formulation contains phyto-estrogens which have pronounced estrogenic activity better than estrogen thus combating the hormone deficiency. It has significant spasmolytic activity mediated through calcium channel blockade. Thus, relieves the pain by direct action on the myometrium. It has proven significant anti-oxidant activity, which by free-radical scavenging, enhances the immunity and general strength of the body. It increases the pain threshold and facilitates better pain tolerance capacity. The formulation has diuretic property which prevents the bloatedness, weight gain and reduces water retention symptoms. The drug alters prostaglandin production by altering arachidonic acid metabolism. Thus, has anti-inflammatory action and relieves the pain. It reduces the acidity of gastric secretions, thus preventing nausea and vomiting. It inhibits platelet aggregation, thus preventing clot formation. It reduces irritation of intestines to cure diarrhoea. - Analysis of the data

of the present study revealed Chaturbeeja *churna* has significant role in the managementof *Kashtartava* (Dysmenorrhoea). Though the results are good, but further study on large numbers of patients should be done along with some specific investigations like Prostaglandin synthetase evaluation. Increasing body of evidences suggests that oxidative stress plays a vital role in the induction and progression of various disorders, the promising antioxidant effect of fenugreek needs to be explored in this area.

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REFERENCES

- 1. Parekh J, Chanda S, In vitro antibacterial activity of the crude methanol extract of *Woodfordia fruticosa* kurz. flower (Lythraceae). Brazilian J Microbiol 2007; 38:204-207
- 2. Harlow S, Park M. A longitudinal study of risk factors for the occurrence, duration and severity of menstrual cramps in a cohort of college women. Br J Obstet Gynaecol 1996; Nov 103(11):1134-42.[Pubmed] PMID:8917003
- 3. Banikarim C, Chacko, M. R. and Kelder, SH, Prevalence and impact of Dysmenorrhoea on Hispanic female adolescents. Archives of Pediatrics & Adolescent Medicine, 2000, Dec;154(12):1226-9.[Pubmed] PMID: 11115307
- 4. Emans SJ, Laufer RL, Goldstein DP The physiology of puberty, Pediatric & adolescent gynecology. 5th ed. Philadelphia. Lippincott, Williams & Wilkins; 2005. p. 417–22.
- 5. Twigg J. Dysmenorrhoea. Curr Obstet Gynaecol 2002; 12:341–45
- 6. Jamieson DJ, Steege JF. The prevalence of dysmenorrhea, dyspareunia, pelvic pain, and irritable bowel syndrome in primary care practices. Obstet Gynecol 1996 Jan;87(1):55–58. [Pubmed] PMID: 8532266
- 7. Ng TP, Tan NC, Wansaicheong GK. A prevalence study of dysmenorrhea in female residents aged 15–54 years in Clementi Town, Singapore. Ann Acad Med Singapore 1992 May;21(3):323–27 [Pubmed] PMID: 1416778
- 8. Andersch B, Milsom I. An epidemiologic study of young women with dysmenorrhea. Am J Obstet Gynecol 1982 Nov 15;144(6):655–60.[Pubmed] PMID: 7137249
- 9. Hirata M, Kumabe K, Inoue Y. Relationship between the frequency of menstrual pain and body weight in female adolescents. Nihon Koshu Eisei Zasshi 2002 June ;49(6):516-24.[Pubmed] PMID: 12138714
- 10. Pawlowski B. Prevalence of menstrual pain in relation to the reproductive life history of women from the Mayan rural community. Ann Hum Biol 2004 Jan-Feb;31(1):1-8.[Pubmed] PMID:14742161
- 11.Pullon S, Reinken J, Sparrow M. Prevalence of dysmenorrhoea in Wellington women. N Z Med J 1988 Feb 10;101(839):52-54.[Pubmed] PMID:3380425
- 12. Strinic T et al Anthropological and clinical characteristics in adolescent women with dysmenorrhea. Coll Antropol 2003Dec ;27(2):707.11.[Pubmed] PMID:14746162
- 13. This and that: the essential pharmacology of herbs and spices. Trends Pharmacol Sci 1992;13:15 20. Bhava Prakasha Nighantu of Sri Bhava Mishra commentary by Dr.K.C. Chunekar edited by Dr.G.S. Pandey published Chaukhambha Bharati Academy Varanasi Reprint 1999. page: 25,32,33,38,39.
- 14 Raja Nighantu by Dr.Indradeva Tripathi published by Chaukhambha Krishnadas Academy Varanasi 4thediton 2006.page:147,148,141.
- 15. Nighantu Adarsha by Bapalal.G.Vaidya Purvardha published by Chaukhambha Bharati Academy Varanasi Reprint year 2007.page:21,74,411,669,679.
- 16. Dhanvantri Nighantu edited by Prof PriyaVrat Sharma published by Chaukhambha Orientalia Varanasi 4nd Edition 2005.page:80,82,86,167.
- 17. Kaiyadeva Nighantu edited by Prof PriyaVrat Sharma published by Chaukhambha Orientalia Varanasi 2nd Edition 2006.page:222,219. Dr.K.M.Nadakarni's Indian Materia Medica published by Popular Prakashan Bombay 1976 vol 1-page 280, 408, 736, 854 & 1240.
- 18. Krithikar.K.R & Basu.B.D, Indian Medicinal Plants, published by International book distributors vol-1 page 174, 700-701,Vol 2:Page:1201,1204.
- 19. Mishra Brahmashankara editor. Bhavaprakasha Nighantu Pur. 3/206. Chaukhamba Sanskrit Sansthan, Varanasi. 1993.
- 20. Fazli FRY, Hardman R. The spice, fenugreek (Trigonella foenum graecum): its commercial varieties of seed as a source of disogenin. Trop Sci 1968;10:
- 21. AL-Habori M, Raman A. Antidiabetic and hypocholesterolaemic effect of fenugreek. Phytother Res 1998;12:233-42.

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- 22. AL-Habori M, Raman A. Pharmacological properties of Trigonella foenum graecum. In: Petropoulos GA, editor. Medicinal and aromatic plants industrial profiles. London7 Taylor & Francis; 2002. p.162–82.
- 23. Blumenthal M, Busse W, Amp R, Goldberg A, The complete commission monograph: Therapeutic guide to herbal medicines, MA: Integrative Communications, Boston; 1988. p.130.
- 24. Hemavathy J, Prabhakar JV, Lipid composition of fenugreek (Trigonella foenum Graecum L.) seeds. Food Chem 1989; 31:1-7.
- 25. Krishnaswamy K, Traditional Indian spices and their health significance, Asia Pac J Clin Nutr 2008;17:265-68.
- 26. Basch E, Ulbricht C, Kuo G, Szapary P, Smith M. Therapeutic applications of fenugreek. Altern Med Rev 2003; 8:20-27.
- 27. Balaraman R, Dangwal S, Mohan M. Antihypertensive effect of *Trigonella foenum-greacum* seeds in experimentally induced hypertension in rats, Pharmaceut Biol 2006;44:568-75
- 28. Mitra A, Bhattacharya DP. Effects of fenugreek in type 2 diabetes and dyslipidaemia, Indian J Practising Doctor 2006;3:14-18.
- 29. Sebastiana KS, Thampanb RV. Differential effects of soybean and fenugreek extracts on the growth of MCF-7 cells. Chemico-Biol Interactions 2007;170:135-43.
- 30. Al-dalain S, El-kutry MS, Ibrahim HS. Inhibitory effect of aqueous extracts of barley and fenugreek on ulcer induction in rats. World App Sci 2008:5:332-39
- 31. Mathern JR, Raatz SK, Thomas W, Slavin JL, Effect of fenugreek fiber on satiety, blood glucose and insulin response and energy intake in obese subjects, Phytother Res 2009;23:1543-48.
- 32. Gupta SK, Kalaiselvan V, Srivastava S, Saxena R, Agrawal SS, *Trigonella foenum-graecum* (fenugreek) protects against selenite-induced oxidative stress in experimental cataractogenesis. Biol Trace Elem Res 2010; Sep; 136(3):258-68. Epub 2009 Oct 13.
- 33. Ruth T. Monograph on Dioscorea Spp. www.phytotherapies.org.
- 34. Chapagain B, Wiesman Z, Variation in diosgenin level in seed kernels among different provenances of *Balanites aegyptiaca* Del (Zygophyllaceae) and its correlation with oil content. Afr J Biotechnol 2005;4:1209-13.
- 35. Duke JA editor. Green Pharmacy, Rodle Press, Pennsylvania 1997;87-90.
- 36. Shukla AS, Chandra S, Bigoniya P. Phytochemical and CNS activity of *Lepidium sativum* Linn. seed's total alkaloid, Scholars Research Library Der Pharmacia Lettre, 2011,3(2): 226-237
- 37. Chopra RN, Chopra IC, Handa KL, Kapur LD. Indigenous Drugs of India, Academic Publishers, Calcutta. 1982, p.58.

- 38. Vohra SB, Khan MSY. Pharmacological Studies on Lapidium Sativum, Linn.Indian J Physiol Pharmacol 1977 Apr Jun; 21(2):118-20.[Pubmed] PMID:885595
- 39. Navarro E, Alonso J, Rodriguez TR, Boada J.Diuretic action of an aqeous extract of Lepidium Latifolium L. J Ethnopharmacol 1994; Jan41(1-2):65-9,[Pubmed] PMID:8170161
- 40. Yadav YC, Jain A, Srivastava DN, Jain A. Fracture healing activity of ethanolic extract of *Lepidium sativum* L. seeds in internally fixed rats' femoral osteotomy model. Int J Pharmacy Pharmace Sci 2011; 3 Suppl 2:193-97.
- 41. Rehman N, Malik HM, Khalid M. Al K, Gilani AH. Studies on antidiarrheal and antispasmodic activities of *Lepidium sativum* crude extract in rats, analgesic activity of the seed of *Lepidium sativum* Linn Ayu. 2010 Jul-Sep; 31(3): 371–373.doi: 10.4103/0974-8520.77163.
- 42. Kaur GJ, Arora DS. Antibacterial and phytochemical screening of *Anethum graveolens, Foeniculum vulgare* and *Trachyspermum ammi*. BMC Complementary & Alternative Medicine 2009; 9-30[Pubmed]. PMID-19656417
- 43. Gilani AH, Jabeen Q, Ghayur MN, Janbaz KH, Akhtar MS. Studies on the antihypertensive, antispasmodic, bronchodilator and hepatoprotective activities of the *Carum copticum* seed extract, Journal of Ethnopharmacology 2005; 98(12):127-35.
- 44. Él-Tahir KEH, Al-Tahir AY, Ageel AM. Recent advances in the pharmacology of the blackseed (*Nigella sativa*). International Symposium on Herbal medicinal plants. Jeddah King Fahad Hospital. 27th March 1997.
- 45. Ashour MM, El-Tahir KEH, Morsi MG, Aba-Alkhail NA. Effect of the volatile oil of *N. sativa* seeds and its components on body temperature of mice. Elucidation of the mechanisms of action. Natr Products Sci 2006; 12: 11-16.
- 46. Sultan MT, Butt MS, Ahmad RS, Pasha I, Ahmad AN, Qayyum MM. Supplem Reducing oxidative stress caused by potassium bromate. entation of *Nigella sativa* fixed and essential oil mediates potassium bromate induced oxidative stress and multiple organ toxicity. Department of Food Sciences, Bahauddin Zakariya University, Multan.[4]Wikipedia,Potasiumbromate http://en.wikipedia.org/wiki/Potassium_bromate).
- 47. Mohamed LS. Immunomodulatory and therapeutic properties of the *Nigella sativa* L. seed. International Immunopharmacology 2005; 5(13–14):1749–70.