## **Review Article**

Kenkyu Journal <u>of Pharmacology</u>



# Antilithiasis, Diuretic, Abortifacient, Antitussive, Nematicidal, Anthelmintic and Antifilarial Seeds used as Therapeutic Agents

Saleemullah Abro<sup>1</sup>, Shah Murad<sup>2</sup>, Wajid Abro<sup>3</sup>, Nusratullah Khan<sup>4\*</sup>, Moosa Khan<sup>5</sup> and Seema Shah Murad<sup>6</sup>

<sup>1</sup>AP Baqai Medical University, Physiology Department, Karachi Pakistan

<sup>2</sup>HOD Pharmacology IMDC and DANTH, Islamabad Pakistan

<sup>3</sup>Microbiology Department at BMSI, JPMC Karachi Pakistan

<sup>4</sup>Assistant Prof of Biochemistry at Bolan University of Medical and health Sciences, Quetta, Pakistan

<sup>5</sup>Chairman Pharmacology at SZABMU Islamabad Pakistan

<sup>6</sup>Gynecologist at NMC, IMDC and DANTH Pakistan

\***Corresponding author:** N Khan, Biochemistry Department at Quetta Institute of Medical Sciences, BMC Quetta, Balochistan, Pakistan. Tel: +923142243415;

Received: December 06, 2019; Accepted: December 13, 2019; Published: December 20, 2019

**Citation:** Saleemullah Abro, Shah Murad, Wajid Abro, Nusratullah Khan, Moosa Khan, et al. (2019) Antilithiasis, Diuretic, Abortifacient, Antitussive, Nematicidal, Anthelmintic and Antifilarial Seeds used as Therapeutic Agents. *Ken Joul Pharcol*, *5*(*1*): *1-3*.

#### Summary

Modest reductions in CHD rates by decreases in saturated fat are possible if saturated fat is replaced by a combination of poly- and mono unsaturated fat, and the benefits of polyunsaturated fat appear strongest. However, little or no benefit is likely if saturated fat is replaced by carbohydrate, but this will in part depend on the form of carbohydrate. Because both N-6 and N-3 polyunsaturated fatty acids are essential and reduce risk of heart disease, the ratio of N-6 to N-3 is not useful and can be misleading. In practice, reducing red meat and dairy products in a food supply and increasing intakes of nuts, fish, soy products and nonhydrogenated vegetable oils will improve the mix of fatty acids and have a markedly beneficial effect on rates of CHD. This study was conducted to see hypolipidemic potential of two medicinal herbs. The research work was single blind placebo-controlled, conducted at Jinnah Hospital, Lahore It was conducted from January 2018 to June 2018. Seventy-five already diagnosed primary and secondary hyperlipidemic patients were selected with age range from 17 to 65 years. Diabetes mellitus, cigarette smoking/alcohol addictive patients, peptic ulcer disease, hypothyroidism, kidney dysfunction, any heart disease and liver disease. All patients were divided in three groups (group-I, group-II, group-III), 25 in each group. All participant's baseline lipid profile data were taken and filed in specifically designed Performa, at start of taking medicine. Twenty-five patients of group-I were advised to take 10 grams of Flaxseeds in three divided doses after meal. Twenty-five patients of group-II were advised to take Ajwain seeds 10 grams in three divided doses after each meal for two months. Twenty-five patients of group-III were provided placebo capsules, (containing grinded rice), taking one capsule after each meal. All participants were advised to take these medicines for eight weeks. Follow-up period: All participants were called fortnightly for their query and follow up. Their LDL-cholesterol, and HDL-cholesterol was determined at the hospital laboratory. In two months, therapy by Flaxseeds decreased LDL-cholesterol from 195.11±2.11 mg/dl to 190.22±3.11 mg/dl, which is significant statistically. HDL was increased from 34.53±1.65 mg/dl to 38.97±2.29 mg/d, which is also significant change. In two months, therapy by AJWAIN, LDL-c reduced from 201.51±2.62 mg/dl to 197.11±2.66 mg/dl, which is significant statistically. HDL-cholesterol increased by Ajwain from 36.97±3.32 mg/dl to 37.45±1.87 mg/dl, which is insignificant statistically. It was concluded from this study that Flaxseeds have more effect on HDL-c but Ajwain has lowest effect on this parameter.

#### Introduction

Meta analysis of previously conducted studies suggests an almost 2.5-fold rise in the prevalence of CAD in two decades from 3.6% in the 1970s to 9.5% in the 1990s in people aged  $\geq$ 35 years in urban Pakistan. With increasing urbanisation in the subcontinent, CAD prevalence can only continue to rise, and this underlines the need to understand risks and determinants of disease in this population to inform preventive strategies. Various studies have reported that unlike European populations, the burden of cardiovascular risk factors in Indo-Asians, such as hypertension, diabetes, dyslipidaemia, obesity, including central obesity, and physical inactivity, are at least equivalent if not greater in women than in men Atherosclerosis is commonly referred to as a "hardening" or furring

of the arteries. It is caused by the formation of multiple atheromatous plaques within the arteries [1]. Flaxseed inhibits the production of pro-inflammatory cytokines, eicosanoids, cytokines and plateletactivating factor derived from arachidonic acid (an omega-6 fatty acid) and thus reduces inflammatory responses. One way that Alpha Linolenic Acid helps the heart is by decreasing the ability of platelets to clump together, a reaction involved in the development of atherosclerosis (hardening of the arteries), it acts as natural aspirin [2]. Flaxseed helps to lower high blood pressure, clears clogged coronaries like a sweeper, lowers high blood cholesterol, bad LDL cholesterol and triglyceride levels and raises good HDL cholesterol. Intake of flaxseeds has also been shown to decrease the ratio of LDL to HDL cholesterol in several human studies and to increase the level of apolipoprotein A1, which is the major protein found in HDL cholesterol. Flaxseeds prevent clot formation in arteries, which may result in strokes, heart attacks and thrombosis. Omega-3 Fatty acids present in Flaxseed appear to enhance the mechanical performance and electrical stability of the heart and to protect against fatal arrhythmias [3-7]. Trachyspermum ammi, has been proven to possess various pharmacological activities like antifungal, antioxidant, antimicrobial, antinociceptive, cytotoxic, hypolipidemic, antihypertensive, antispasmodic, broncho-dilating actions, antilithiasis, diuretic, abortifacient, antitussive, nematicidal, anthelmintic and antifilarial. Further, studies reveal the presence of various phytochemical constituents mainly carbohydrates, glycosides, saponins, phenolic compounds, volatile oil (thymol,  $\gamma$ -terpinene, para-cymene, and  $\alpha$ - and  $\beta$ -pinene), protein, fat, fiber and mineral matter containing calcium, phosphorous, iron and nicotinic acid. These studies reveal that T. ammi is a source of medicinally active compounds and have various pharmacological effects; hence, it is encouraging to find its new therapeutic uses [7-9]. The constituents of the seed of Ajwain included carbohydrates (38.6%), fat (18.1%), protein (15.4%), fiber (11.9%), tannins, glycosides, moisture (8.9%), saponins, flavone, and mineral matter (7.1%) containing calcium, phosphorous, iron, cobalt, copper, iodine, manganese, thiamine, riboflavin, and nicotinic acid [10,11]. Antiplatelet-aggregatory experiments in vitro with blood from human volunteers, it that a dried ethereal extract of Ajwain seeds, inhibited aggregation of platelets induced by arachidonic acid, collagen and epinephrine [12]. Antihyperlipidemic effect of Ajwain seed has been proved by researchers [13].

five already diagnosed primary and secondary hyperlipidemic patients were selected with age range from 17 to 65 years. Exclusion criteria were, diabetes mellitus, cigarette smoking/alcohol addictive patients, peptic ulcer disease, hypothyroidism, kidney dysfunction, any heart disease and liver disease. All patients were divided in three groups (group-I, group-II, group-III), 25 in each group. Their baseline lipid profile data was taken and filed in specifically designed Performa, at start of taking medicine. Twenty-five patients of group-I were advised to take 10 grams of Flaxseeds in three divided doses after meal. Twenty-five patients of group-II were advised to take Ajwain seeds 10 grams in three divided doses after each meal for two months. Twenty-five patients of group-III were provided placebo capsules, (containing grinded rice), taking one capsule after each meal. All participants were advised to take these medicines for eight weeks. All participants were called fortnightly for their query and follow up. Their LDL-cholesterol, and HDLcholesterol was determined at the hospital laboratory. After two months therapy results were compared and data were expressed as the mean ± Standard Deviation and 't' test was applied to determine statistical significance as the difference. A probability value of <0.05 was considered as non-significant and P<0.01 was considered as significant change in the results when pre and post-treatment results were compared.

### Results

When results were compiled and statistically analyzed it was observed that Flaxseeds and Ajwain decreased LDL-cholesterol, and increased HDL-cholesterol significantly as compared to placebo therapy. Before treatment and after treatment values and results are shown in table 1, 2 and 3.

#### Patients & Method

The research work was single blind placebo-controlled, conducted at Jinnah Hospital, Lahore from January 2018 to June 2018. Seventy-

 Table 1: Showing Effects of Placebo Therapy Before and After Treatment with Its Statistical Significance in Group-III Patients (n=25)

LDL-c	188.11±1.06	187.77±2.51	0.3	>0.05
HDL-c	30.78±2.65	31.39±1.66	0.6	>0.05

#### Table 2: Showing Effects of Flaxseeds Before and After Treatment with Its Statistical Significance in Group-I Patients (n=22)

Parameter	At start	At end	Diff	p-value
LDL-c	195.11±2.11	190.22±3.11	4.9	< 0.01
HDL-c	34.53±1.65	38.97±2.29	4.4	< 0.01

#### Table 3: Showing Effects of Flaxseeds Before and After Treatment with Its Statistical Significance in Group-II Patients (n=24)

LDL-c	201.51±2.62	197.11±2.66	4.4	< 0.01
HDL-c	36.97±3.32	37.45±1.87	0.5	>0.05

**KEY:** All values are measured in mg/dl. LDL-c= low density lipoprotein cholesterol, HDL-c= high density lipoprotein cholesterol. P-value <0.01 stands for significant change, P-value <0.05 stands for non-significant change. n stands for sample size.

#### Discussion

The greater prevalence of definite CAD in men is largely due to the effect of smoking exposure, which owing to its substantially higher prevalence in men, overwhelms the impact of the greater prevalence of the metabolic syndrome in women [14]. Angina is the most common symptom of CAD. Flaxseeds and Ajwain are thought to inhibit these pro-inflammatory effects in human body. In our results using 10 grams of Ajwain by 24 hyperlipidemic patients for the period of two months, LDL-c reduced from 201.51 $\pm$ 2.62 mg/dl to 197.11 $\pm$ 2.66 mg/dl. Difference in pre and post treatment values is 4.4 mg/dl. Increase in HDL was 0.5 mg/dl, which is non-significant change in pre and post treatment values. In our study Flaxseeds decreased LDL-c from 195.11 $\pm$ 2.11 to 190.22 $\pm$ 3.11 mg/dl in two months therapy by 10 grams Flaxseeds used by 22 hyperlipidemic patients [15,16]. Difference in pre and post treatment values is 4.9 mg/dl in this parameter. HDL was increased from 34.53 $\pm$ 1.65 to 38.97 $\pm$ 2.29 mg/dl. Difference in precentage when measured/calculated it was 4.4 mg/dl which is significant bio statistically with p-value <0.01. These results match with results of study conducted by Cho Y et al [17]. who proved almost same effects on two lipid profile parameters ie; LDL-cholesterol and HDL-cholesterol. Chaudhury S [18]. Proved same changes in LDL-cholesterol and HDL-cholesterol which also support our results biostatistical. Nagalakshmi S et al [19]. Explained that all herbs with their therapeutically medicinal potential will work when used in high amount and for long period. Jenkins D et al [20]. Described that Flaxseeds or its oil preparation have same effects on all parameters of lipid profile. On comparison between statins and herbal medicine having hypolipidemic effects. Kelley DS et al [21]. Explained that there

is too much difference in hypolipidemic effects of allopathic medication and herbs, having less potent hypolipidemic features of herbal medications. Shahidi F and Miraliakba H [22]. Proved that all parameters of lipid profile including total, LDL-cholesterol, HDL-cholesterol, VLDL-cholesterol, IDL and triglycerides are affected by Flaxseeds oil preparations. They have focused on inhibition of enterohepatic circulation of bile acids and explained that due to lack of bile acid pool in gall bladder, hepatocytes start to synthesize bile acids instead of cholesterol synthesis. Many scientists proved that if used Flaxseeds with dietary restrictions and change in sedentary life style, HDL-cholesterol will increase surely. They explained that if only one parameter of lipid profile ie; HDL-cholesterol is increased, all other parameters in ratio will obviously be reduced leading to lesser chances of development of CAD. According to some scientist's anti-inflammatory effects of Flaxseeds play key role in prevention of atherosclerosis and CAD. If sedentary life style is changed by hyperlipidemic patients, very small but regular amount of Flaxseeds are required to stay at preventive step of coronary artery disease due to Hyperlipidemia. High LDL-cholesterol has close concerned with pro-inflammatory responses leading to platelet aggregation [1,8,13]. There's a type of hyperlipidemia that you can inherit from your parents or grandparents. It's called familial combined hyperlipidemia. Familial combined hyperlipidemia causes high cholesterol and high triglycerides. People with this condition often develop high cholesterol or high triglyceride levels in their teens and receive a diagnosis in their 20s or 30s [23-27].

#### References

- 1. Genwae E, Reth T, Manhas A, Farmer JA (2016) Hypolipidemic therapy and cholesterol absorption. CAR 11: 88-94.
- 2. Meth G, Chialva F, Monguzzi F, Manitto P, Akgül A (2017) Essential oil constituents of Trachyspermum copticum (L.) Link fruits. JEOR 12: 90-93.
- Herfg G, Nagalakshmi S, Shankaracharya NB, Naik JP, Rao LJM (2018) Studies on chemical and technological aspects of ajowan (Trachyspermum ammi syn. Carum copticum) JCMR 111: 1110-1117.
- 4. Srivastava KC (2011) Extract of a spice-omum (Trachyspermum ammi shows antiaggregatory effects and alters arachidonic acid metabolism in human platelets. Prostaglandins Leukot Essent Fatty Acids 33: 16.
- 5. Anilakumar KR, Saritha V, Khanum F, Bawa AS (2009) Ameliorative effect of ajwain extract on hexachlorocyclohexane-induced lipid peroxidation. Food Chem Toxicol. 47: 279-282.
- 6. Chialva F, Monguzzi F, Manitto P, Akgül A (2012) Essential oil constituents of Trachyspermum copticum (L.) and Lipid Metabolism. J Essent Oil Res 5: 105-106.
- Singh G, Maurya S, Catalan C (2009) Chemical, antifungal, antioxidative studies of Ajwain oil and its acetone extract. J Agric Food Chem 52: 3292-3296.
- Choudhury S (2013) Composition of the seed oil of Trachyspermum ammi (L.) Sprague from northeast India. J Essent Oil Res 10:588-590.
- 9. Murthy PS, Borse BB, Khanum H, Srinivas P (2009) Inhibitory effects of Ajwain (Trachyspermum ammi) ethanolic extract on A. ochraceus growth and ochratoxin production. Turk J Biol 33: 211-217.
- Patenaude A, Rodriguez-Leyva D, Edel A L, Dibrov E, et al. (2009) Bioavailability of a-linolenic acid from flaxseed diets as a function of the age of the subject. Euro. J. Clin. Nutri 63: 1123-1129.
- 11. Krishnamoorthy V, Madalageri MB (2010) Bishop weeds (Trachyspermum ammi): An essential crop for north Karnatka. J Med Aromat Plant Sci 21: 996-998.
- 12. Ishikawah T, Sega Y, Kitajima J (2011) Water-soluble

constituents of ajowan. Chem Pharm Bull 49: 840-844.

- Rodriguez-Leyva D, Bassett C M, Mccullough R, Pierce G N (2010) The cardiovascular effects of flaxseed and its omega-3 fatty acid, alpha-linolenic acid. Can. J. Cardiol. 26: 489-496.
- Tzang B S, Yang S F, Fu S G, Yang H C (2009) Effects of dietary flaxseed oil on cholesterol metabolism. Food Chem 114: 1450-1455.
- 15. Prasad K (2005) Hypocholesterolemic and antiatherosclerotic effect of flax lignan complex isolated from flaxseed. Atherosclerosis 179: 269-275.
- Arjmandi B, Khan D, Juma S (2011) Whole flaxseed consumption lowers serum LDL-cholesterol and lipoprotein (a) concentrations in postmenopausal women. Nutrition Research 18: 1203-1214.
- 17. Cho Y, Kwon E, Kim H (2009) Low trans structured fat from flaxseed oil improves plasma and hepatic lipid metabolism. Food and Chemical Toxicology 47:1550-1555.
- Choudhury S (2011) Shankaracharya NB, Naik JP, Rao LJM Composition of the seed oil of Trachyspermum ammi (L.) Sprague from northeast India. J Essent Oil Res 10: 588-590.
- Nagalakshmi S, Shankaracharya NB, Naik JP, Rao LJM (2011) Studies on chemical and technological aspects of ajowan (Trachyspermum ammi syn. Carum copticum) J Food Sci Technol 37: 277-281.
- 20. Jenkins D, Kendall C, Shankaracharya NB, Naik JP, Rao LJM Vidgen E, et al. (2010) Health aspects of partially defatted flaxseed, including effects on serum lipids, and oxidative stress: a controlled crossover trial. American Journal of Clinical Nutrition 69: 395.
- Kelley D S, Siegel D, Vemuri M, Mackey B E (2007) Docosahexaenoic acid supplementation improves fasting and postprandial lipid profiles in hypertriglyceridemic men. Am. J. Clin. Nutr 86: 324-333.
- Shahidi F, Miraliakbari H, Shankaracharya NB, Naik JP, Rao LJM (2004) Omega-3 fatty acids in health and disease: Part1cardiovascular disease and cancer. J. Med. Food 7: 387-401.
- 23. Bert VV, Shankaracharya NB, Naik JP, Rao LJM Anilakumar KR, Saritha V, et al. (2017) Ameliorative effect of ajwain extract on hexachlorocyclohexane-induced lipid peroxidation. JFB 13: 56-59.
- Jhialva F, Shankaracharya NB, Naik JP, Rao LJM Monguzzi F, Manitto P, et al. (2012) Essential oil constituents of Trachyspermum copticum (L.) and Lipid Metabolism. J Essent Oil Res 5: 105-106.
- 25. Lingh G, Maurya S, Shankaracharya NB, Naik JP, Rao LJM Catalan C (2009) Chemical, antifungal, antioxidative studies of Ajwain oil and its acetone extract. J Agric Food Chem 52: 3292-3296.
- Ghoudhury S (2013) Shankaracharya NB, Naik JP, Rao LJM Composition of the seed oil of Trachyspermum ammi (L.) Sprague from northeast India. J Essent Oil Res 10: 588-590.
- Furthy PS, Shankaracharya NB, Naik JP, Rao LJM Borse BB, Khanum H, et al. (2009) Inhibitory effects of Ajwain (Trachyspermum ammi) ethanolic extract on A. ochraceus growth and ochratoxin production. Turk J Biol 33: 211-217.

**Copyright:** ©2019 N Khan, Biochemistry, et al. This is an open-access article distributed under the terms of the Creative Commons Attribution License, which permits unrestricted use, distribution, and reproduction in any medium, provided the original author and source are credited.