# Hypolipidemic Dietary Components

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# ABSTRACT

Several prospective epidemiological studies over the past 20 years concluded that ingestion of certain food or dietary components improves serum lipid profile and are effective in management of hyperlipidemia. The objective of this paper is to identify and quantify selective lipid lowering dietary components. This review focuses on identifying, quantifying and understanding the possible mechanism of action of soy, flaxseeds, oats and psyllium in improving lipid profile and reducing atherosclerotic cardiovascular disease (ASCVD) risk factor. A systematic search of database was performed to retrieve studies of published human studies and trials from 1995-2012, concentrating on recent systematic reviews, meta-analysis and intervention trials done to further elucidate the role of selective food (*soy, oats, flaxseeds and psyllium*) and their bioactive components (*soy protein, soy isoflavones, flaxseed lignan, flaxseed onega 3 fatty acid, b-glucan, insoluble fiber*) involved in improving lipid profile and reducing the cardiovascular disease risks. The findings quantify the amounts showing beneficial effects on serum lipid profile. Overall, the review of these clinical evidences suggested that inclusion of these food items can moderately but significantly reduce the risk of dyslipidemia, when consumed regularly as a part of a healthy diet and thus may help reduce any untoward cardiovascular event

Key word: Soy, Flaxseeds, Psyllium, Oats, and lipid profile.

# INTRODUCTION

Cardiovascular diseases (CVD) are the most prevalent cause of death and disability in both developed as well as developing countries. INDIA is one of the leading nations in ASCVD and is already a global capital of diabetes. In 2005, 29% of deaths in INDIA occurred alone due to CVD<sup>1</sup>. According to the recent estimates about 2.9 crores India's population is burdened with CVD and may increase to 6.4 crores, if no early intervention steps are taken. Dyslipidemia has been found to be one of the most important contributing factors<sup>2</sup>.

Dyslipidemia describes a number of abnormalities in lipoprotein homeostasis including hypercholesterolemia and hypertriglceridemia. Several co-morbid conditions like myocardial infarction, stroke, atherosclerosis, nephropathy, fatty liver cirrhosis and many others are associated with dyslipidemia. Therefore, reductions in dyslipidemic conditions and eventually CHD risk begin with the adoption of a healthy lifestyle. Thus, as a part of a comprehensive strategy to reduce complications of hypercholesterolemia, NCEP (*National Cholesterol Education Programme*) endorsed the use of several lipid lowering dietary agents<sup>3</sup>.

This review emphasize on the quantity of soy, flaxseeds, oats and psyllium that have shown to improve the lipid profile, the mechanisms of each bioactive component responsible in reducing the risk of ASCVD and several human clinical trials data regarding their safety and efficacy.

#### **Bioactive components**

Dietary Fiber (DF) are the edible parts of plants or analogous carbohydrate that are resistant to digestion & absorption in the human intestine with complete or partial fermentation in the large intestine and which promote beneficial physiological effects including laxation and/or blood glucose attenuation and /or blood cholesterol attenuation.

Total dietary fiber (TDF) occurs in 2 major forms i.e. soluble dietary fiber (SDF) and insoluble dietary fiber (IDF). (4.) DF content of various foods are mentioned in Table1

# **Mechanism of Action**

Soluble fiber lowers total and LDL-c cholesterol which can be attributed to enhanced gastric emptying, intestinal binding of bile acids thus reducing entero-hepatic bile recirculation, lowered dietary cholesterol absorption, affects VLDL composition which then contains more amount of phospholipids and less of cholesterol, also improves satiety. It also undergoes fermentation to short chain fatty acids (SCFA) in the colon thus reducing endogenous synthesis of cholesterol. It also decreases dietary glucose absorption thereby down regulating cholesterol synthesizing anabolic hormone insulin release<sup>7-13</sup>.

#### Omega-3 fatty acid from a plant source

They are essential fatty acids for competitive synthesis of eicosonoids which are antiinflammatory, anti aggregatory and anti-thrombotic in action<sup>14-15</sup>. Additional physiologic functions of omega 3 fatty acids which may prevent ASCVD include several mechanisms such as; maintain fluidity of the cell membrane thus facilitating removal of cholesterol, down regulating phosphotidic acid phosphatase (PAP) and diacylglycerol acyl tranferase (DGAT) thus reducing fatty acid synthesis, also activates PPAR involved in B-oxidation of fatty acids, upregualting lipoprotein lipase (LPL) activity thus increasing fats hydrolysis. N-3 fatty acids also modulate sterol regulating element binding protein (SREBP) and increases degradation of apoprotein B which eventually decreases LDL-c secretion and transport<sup>16-18</sup>. Table 2 illustrates nutrient composition of flaxseed

# Phytoestrogens

Are a diverse group of naturally occurring non steroidal plant compounds that, because of their structural similarity with estradiol (17- $\beta$ estradiol), have the ability to cause estrogenic or/ and antiestrogenic effects<sup>20</sup>. Phytoestrogen content of selective foods are discussed in Table 3. There are 2 major classes of phytoestrogens including: Isoflavones contains major bioactive components i.e. daidzein and genistein. They are primarily present in soy as much as 3 mg/g dry weight of soy. Lignans and their major bioactive components secoisolariciresinol and matairesinol are primarily found in flaxseeds<sup>21</sup>.

# **Mechanism of Action**

Clinical evidence suggest that phytoestrogens is effective in lowering serum cholesterol via a series of mechanism which include; up regulation of LDL-c receptor activity thus increasing its clearance, it also depresses the activity of platelet activating factor receptor antagonists eventually reducing oxidation of the lipid, the primary step in process of hypercholesterolemic atherosclerosis. Phytoestrogens also enhances the activity of cholesterol-7 hydoxylase which in turn decreases endogenous cholesterol synthesis. There is also down regulation of the enzyme tyrosine kinase thus reducing thrombin production and ultimately myocardial infarction risk (MI) risk<sup>22-27</sup>.

# Soy protein

Recently soybeans are considered by many agencies as a source of a complete protein. Hence variety of foods such as salad dressings, beverage powders, cheese, infant formulas and others contains soy proteins. The nutritional value of soy protein is equivalent to that of animal protein of high BV. For instance, isolated soy protein has a PDCAAS (Protein Digestibility Corrected Amino Acid Score) of 1.0, which is the same as that of casein and egg protein. 100g of Whole soybeans contains 43.2 g of protein<sup>28-30</sup>. Biological value of various soy products like whole soybean, soy milk, soy protein isolates are 96.0, 91.0, and 74.0 respectively<sup>31</sup>.

# **Mechanism of Action**

Various studies have shown that soy proteins effectively lowers cholesterol through series of mechanisms such as; it binds bile acids thus reducing enterohepatic recirculation of bile, it also up regulates the apo B and apo E receptor activity thus reduces serum LDL-c levels. Soy proteins have also shown to reduce HMG-CoA reductase activity, the rate limiting enzyme in cholesterol synthesis and increase cholesterol-7 hydroxylase activity which eventually lowers cholesterol synthesis<sup>31-33</sup>.

# **Human studies**

To identify the majority of human studies on cardiovascular effects of these food items we performed a systematic search of the following

	TDF (g)/100g	IDF (g)/100g	SDF (g)/100g
Wheat	12.5	9.6	2.9
Oat Bran	5.0	-5.0	
Oats, Whole (1/2 Cup Cooked)	1.6	1.1	0.5
Soybean	23	17.9	5.1
Black Gram, Whole	20.3	15.4	4.9
Bengal Gram ,Whole	28.3	25.2	3.1
Flax Seeds	27.3	22.0	5.3
Gingelly Seeds	16.8	13.6	3.2
Psyllium Husk (10 g)	8	0.9	7.1
Apple	3.2	2.3	.9
Guava	8.5	7.1	1.4
Sapota	10.9	9.1	1.8
Carrot	4.4	3	1.4
Peas, Green	8.6	7.2	1.4

Table 1: Dietary fiber content of	selected	foods <sup>5,6</sup>
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(TDF: Total Dietary Fiber; IDF: Insoluble dietary fiber; SDF: soluble dietary fiber)

	Tab	le 2: Various	Flaxseed P	roducts Composi	tion	
Form of flaxseeds	Weight (g)	Energy (kcal)	ALA (g)	Total dietary fiber(g)	Soluble fiber (g)*	SDG content (mg)*
Whole seed	11	50	2.5	3.0	0.75	8.8
Ground seeds	8	36	1.8	2.2	0.55	6.4

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ALA=Á Linolenic Acid, SDG: Secoisolariciresinol Diglucoside

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Table 3: Phytoestrogens content of foods <sup>21</sup>	Table 3:	Phytoestrogens	content	of	foods <sup>21</sup>
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Flaxseed oil

Phytoestrogens content	(µg/100g)
Flax seed	379380
Soy beans	103920
Tofu	27150.1
Soy yogurt	10275
Sesame seed	8008.1
Flax bread	7540
Soy milk	2957.2
Dried dates	329.5

databases: PubMed, Medscape, MDconsult etc from 1995-2009. We used the headings "soluble fiber, botanical n-3 fatty acids, Phytoestrogens and soy proteins." and searched the terms "soy, flaxseeds, oats and psyllium." using this strategy we identified 75 relevant articles and book chapters. The description below highlights the information on potential CV application of this food items.

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# **Epidemiological data**

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Epidemiological data concerning intake of fiber, n-3fatty acids, phytoestrogens and soy

Rafaranc a	Study design	Sample size	Intervent ion	Duration	Control group	Change in TC	Change in LDL	change in HCL	change in TG	Other CV effects
Pipe E, et al; 2009. (36.)	Double blind, RCT, placebo control	29 type 2 diabetic	SPI containin g 80mg iso1avon es	57 days	Normal diet without SPI	ιP=<0.00 2	ιP=<0.04	ι P=<0.02	No change	Decrease s apoBapo A p=<0.05
Taku K, et al; 2007. (37.)	RCT	11 trials from 1990- 2006	Iso <b>ta</b> von es=64- 318mg/d Soy proteins= 25- 133g/d	Meta- analysis	No control	ι <b>Ρ =</b> 0.02	ιΡ < 0.0001	יP = 0.05	No change	Significan t in hyperchol esterole mic subjects
Clerici CC, et al. ; 2007. (38.)	Uncontrol led trial	62 hyperchol esterole mic	33mg of iso <b>1</b> avon es	×	No control	เ73% (P = 0.001)	ι8.6% (Ρ = 0.002)	No chang	No change	-
Desroche s S., et al; 2004. (39)	RCT	36 mild- moderate hyperchol esterole mic	17g soy protein	6weeks	Animal & soy proteins with,but isoflavon es					Larger LDL peak size, decrease cholester ol levels in small LDL.
Wangen KE, et al.; 2001. (40.)	RCT	18 postmen opausal women	132±22 mg/d Iso <b>1</b> avon es	93d	Control diet	-	(6.5% p=<0.02			ιLDL:HD L by 7.7% p=<0.02
Wofford MR et al. (2012) (41.)	randomiz ed, double- blind, 3- phase crossover trial	352 aduits with serum cholester ol <240mg/ dl	40g soy protein/d	8weeks	Milk protein or complex carbohyd rate from wheat	Compare d with carbohyd rate, soy protein( (P=0.03)	No effect	Compare d with milk protein, soy protein suppleme ntation, HDL( (P %00009)	No effect	l TC:HDL (P=0.03) and (P <0.001) compare d to carbohyd rate and mik protein suppleme ntation respectiv ely

Table 4: Interventional trial data on lipid lowering effects of soy

(RCT-randomized control trial, TC- Total cholesterol, LDL-C- low density lipoprotein, HDL- high density lipoprotein, TG-triglycerides; ↓: Reduced; SPI: Soy protein isolate; P<0.05: Statistically significant)

Refere nce	Study design	Sample size	Interventi on	Durat ion	Compa rator group	dhan ge in TC	Chang e in LDL	dha nge in HDL	cha nge in TG	Other CV effects
An Pan, et al; 2009. (43.)	Meta- analysi s	28 studies from Jan 1990 Oct2008	Whole/def atted/ ground flaxseeds doses ranges from 20.0 to 50.0 g		-	10.1 0 mmo VL	10.08 mmol L			Effects were significant in postmenopau sal and hypercholest erolemic subjects
Patade A. et al; 2008. (44.)	RCT	55 mild- moderate Hypercholest erolemic women	20g whole flaxseeds	3 mont hs	Control	Į7 %	<b>↓10%</b>		-	
Mandaş escu S et al; 2005. (45.)	RCT	40 hyperlipidemi c patients	20 g flax group	6mon ths	Statin	↓17. 2%,	13'8%		(- 36.3 %)	↓TC/HDL-C ratio (- 33.5%).
Jenkins D ,et al.; 2001. (46.)	RCT	29 hyperlipidemi c	≈50 g partially defatted flaxseed/ d	3-mk	whe <i>a</i> t bran	↓4.6 ± 1.2% : P = 0.00 1	↓7.6 ± 1.8%; P < 0.001	-	-	↓apolipoprote in B (5.4 ± 1.4%; P = 0.001), and apolipoprotei n A I (5.8 ± 1.9%; P = 0.005)
B Arjman di , et al; 2000. (47.)	double blind cross- over study	38 mild, moderate, or severely (5.85–9.05 mmo/L) hypercholeste rolemic postmenopau sal women	38 g	6-mk	Sunflow er seeds	169 %	↓p <0.0 O1) Iower LD L- choles terol (14.7 %	-	-	lipoprotein(a) (Lp(a)], significantly (p<0.05) lowered
Fukumit su Set al. 2010 (48.)	double blinded, random ized, and placebo - controll ed study	moderately hypercholeste rolemic men (n=30)	Group 1: flaxseed lignan capsule (20mg/d) Group2: flaxseed lignan capsule (100mg/d )	12 week s	Placebo capsule s					Group 2 v/s control ↓ in LDL/HDL ( P=<0.05)

# Table 5: Interventional trial data on lipid lowering effects of flaxseeds

(RCT-randomized control trial, TC- Total cholesterol, LDL-C- low density lipoprotein, HDL- high density lipoprotein, TG-triglycerides;  $\downarrow$ : Reduced; P<0.05: Statistically significant)

proteins are studied as all the discussed food items are a good sources of this bioactive components. Epidemiological data on dietary fiber and risk of ASCVD are extensive and suggest in general an inverse correlation between them. Several observational trials have found positive correlation between soy proteins and n-3 fatty acids and secondary prevention of ASCVD. Also observational studies revealed that lignans may reduce ASCVD morbidity and mortality.

The following section describes data from human studies that addresses the *Quantity and Effects* of soy, flax, oats and psyllium and their bioactive components on CV risk factors dyslipidemia and others like inflammation, blood pressure and glycemic control.

# Bioactive Components Enriched Food Items Soy

Soy (glycine max) is a species of legume native to East Asia. The beans contain significant amounts of proteins (38%), isoflavones, dietary fiber (25-30%) and phytic acid. The principle carbohydrates (30-32%) of mature soybeans are the disaccharide sucrose (2.5-8.2%), trisaccharde raffinose (0.1-1.0%) and the tetrasaccharide stachyose (1.4-4.1%). The majority of soybean carbohydrates can be classed as belonging to dietary fiber. The Food Drug and Administration (FDA) have approved soy as an official cholesterollowering food, along with other heart and health benefits<sup>35</sup>.

The major bioactive components that are responsible for improving the lipid profile on soy interventions are dietary fiber, phytoestrogens and soy proteins and articles related to these are discussed in table 4.

# To Summarize

The bulk of the evidence from five clinical trials<sup>36-41</sup>. Suggest that soy proteins (20-40g/d) or soy isoflavones (30-150mg/d) can modestly but significantly (p=<0.05) reduce total and LDL cholesterol in both normal and hypercholesterolemic subjects, without a significant effects on HDL-c and TG's. Effects were more prominent in post menopausal women or in subjects with initial mild to moderate hypercholesterolemia.

Desroches S., *et al.*, concluded that soy was also effective in shifting LDL particle size to a less atherogenic pattern. Thus, replacing foods high in saturated fats, trans fats & cholesterol; by soy products containing 30-150 mg/d of isoflavones or 20-40g/d of soy protein have a positive effects against coronary risk factors and related co morbidities e.g.: type2 DM ,hypertension and others.

# Flaxseeds

Flaxseed (linseed) is a smooth, flat and reddish-brown in color is native to the region extending from the eastern Mediterranean to INDIA. Whole flaxseeds contain 28% dietary fiber (7-10% soluble fiber, 11-18% insoluble fiber); 40% fats (57% of omega 3 fatty acids) and 21% proteins. it is also the richest source of phtoestrogens- lignans<sup>42</sup>.

Consumption of flaxseeds have shown to reduce total and LDL cholesterol as well as platelet aggregation<sup>43,44</sup>. The major bioactive components responsible for hypolipidemidic action of flaxseeds are dietary fiber, omega3 fatty acids and lignans and human trials related to these are discussed in detailed in table 5.

# **To Summarize**

The results of the above discussed 5 clinical trials43-48 suggest that flaxseeds (20-50g/d whole or partially defatted respectively) are efficacious in improving the lipid profile in both normal and mild-moderate hypercholesterolemic subjects. This can be due to the bioactive components in flax like fiber, n-3 fatty acids and lignans which via several mechanisms as discussed earlier have shown to have a positive association with reductions in serum total and LDL cholesterol. The effects of flax seeds were more prominent in subjects with initial hypercholesterolemia like in postmenopausal women. They have also shown to have beneficial effects on other CVD risk factors like ratios of TC: HDL, LDL: HDL reductions in Apo B 100 and also Lp (A) a strong predictor of CVD. Thus daily incorporation of 20-50g of whole or partially defatted seeds respectively is effective in improving lipid profile.

# Oats

Oats most commonly are available as

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Referen ce	Study de≋gn	Sample size	Interven tion	Duratio n	Compar ator group	change in TC	Change in LDL	change in HCL	change in TG	Other CV effects
VIIIasmil R, et al.; 2007. (50.)	RCT	38mild- mod. Hyperch olesterol emic male	6g b-glucan	8 weeks	Control	ιΡ=<0.0 01	ιΡ<0.00 1	r27.8% p=≺0.00 1		(TC:HD L (P=<0.0 01) LDL:HD L (P<0.003
Saltzma n et al.; 2001. (51.)	RCT	43adults	45g oats/d	8 weeks	Control diet	ιP=<0.0 03	ι P=<0.0 08	-	-	) Decreas ed SBP (P= 0.026)
Romero A et al; 1999. (52.)	RCT	36 normal 30 hyperch ole <i>s</i> terol emic	2.6g oats fiber & 1.3g psyllium fiber	8 weeks	Control wheat diet		ι26% in oarts 22.6% in psyllium (P= <0.01)		ι28%(P= <0.001)	0.020)
Queene n KM et al.; 2007. (53.)	RCT	75 hyperch olesterol emic subjects	6g beta glucan,4d	6 weeks	Control	ι(0.3mol/ L from baseline	(0.3 mmolΔ from baseline	-	-	Highest SCFA producti on than inulin and guar guar gum, in vivo.
Robitalle a J et al.; 2005. (54.)	RCT	34 premeno pausal	28g oat bran <i>i</i> d	4 weeks	-			(11.2%) (p=0.002 )		ιΤC:HD L by 7.0% (p= 0.002)
Maki K et al. 2010 (55.)	Randomi zed, parallel- arm, controlle d trial.	N=144 Free- living, overweig ht and obese adults with baseline LDL cholester ol levels 130 to 200 mg/dL	3g,kd oats beta- glucan in form of RTE cereal, as a part of reduced energy dietary program	12 weeks	Control RTE cereal	ι₽Φ.03 8	ι₽<0.00 5	NS	NS	Non HDL (P<004 6 Waist circumfe rence (P<001 2
Charfton K et al. 2012 (56.)	RCT parallel singles blind trial	N=87 mildly hyperch olesterol emic men and women	High dose: 3g,Xd beta glucan glucan in form of oats porridge	6weeks	Minimal b glucan	-	l P=0.04 in both high dose and low dose beta	-	-	

# Table 6: Interventional trial data on lipid lowering effects of oats

(RCT-randomized control trial, TC- Total cholesterol, LDL-C- low density lipoprotein, HDL- high density lipoprotein, TG-triglycerides; RTE:Ready-To-Eat ;SCFA: Short chain fatty acid;  $\downarrow$ : Reduced; P<0.05: Statistically significant)

rolled, crushed into oatmeal or ground into oat flour. Oat bran is the outer casing of the oats. Oats contain more soluble fiber as compare to any other grain. The soluble fiber in oats comprises of beta glucan, a class of polysaccharide having mixed linkages. The percentages of beta glucan in various products of whole oats are: oat bran >5.5-23%, rolled oats and oat flour about 4 %. FDA in 1998 approved to claim food products containing 3g of soluble fiber from oats can be labeled as foods reducing risk of

Refere nce	Study de sign	Sample size	Interve ntion	Duratio n	Comparat or group	dhange in TC	Chan ge in LCL	chan ge in HDL	cha nge in TG	Other CV effects
Ganji V, et al.; 2008. (58.)	RCT	11 hypercholeste rolemic postmenopau sal and 8 premenopaus al women	15g psyllium /d	6 weeks		ι52% (p=<0.05) in post menopau sal	-	-		
Sola R et al.; 2007. (59.)	RCT	28 hypercholeste rolernic men	10.5g/d psyllium husk	8 weeks	10.5g psyllium seeds		-	(6.7% p=<0. 006 in husk group	(6.7 % D2 in husk grou P	apoB100:apoA1 =4.7 %,TC:HDL =10.6 %, LDL:HDL=14.2 % in HUSK group
Moreyr a A et al.; 2005. (60.)	Double blind placebo control	68 mild- moderate hypercholeste rolemic	15gpsyll ium +10mg simavas tatin	12 weeks	20mg simvastatin , 10mg simvastatin +placebo		(63m g/dl (p=0. 03) in psylliu m group	-		15g psyllium was as effective as 10mg of simvastatin alone
Ander son JW et al.; 2000. (61.)	Multicent ered sudy	38 primary hypercholeste rolemic men and women	5.1g psyllium twice a day	26week		ι <b>4.7%</b>	ί6.7% p=<0. 001			
Santor e G et al. 2009 (62.)		40 T2 DM on OADA's and controlled diet	3.5g psyllium thrice a day	8 weeks	OADA's + Controlled diet				ιΡ< 0	

Table 7: Interventional trial data on lipid lowering effects of psyllium
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(RCT-randomized control trial, TC- Total cholesterol, LDL-C- low density lipoprotein, HDL- high density lipoprotein, TG-triglycerides; OADA's: Oral anti-diabetic agents; T2DM: Type 2 Diabetes Mellitus; ↓: Reduced; P<0.05: Statistically significant)

heart disease<sup>49</sup>. The major bioactive components in oats responsible for lipid lowering seem to be beta glucan as reported by various trials discussed in table 6.

#### **To Summarize**

The results from these studies<sup>50-56</sup> suggest that oats beta glucan is efficacious in significantly improving lipid profile as it was able to reduce TC and LDL-c in normal as well as hypercholesterolemic subjects significantly (p=< 0.05). Its intervention also significantly increased HDL levels (Villasmil R et al.). It was also effective in decreasing LDL: HDL and TC: HDL ratios from the baseline significantly. Queenen et al., also found that oats beta glucan is a most fermentable fiber and produces the highest levels of SCFA butyrate compared to inulin and guar gum in vivo. Oat bran also proven to be beneficial in premenopausal women as it increased HDL-c levels and reduced LDL and TC levels Oats was also able to reduce classical CVD risk marker i.e. systolic blood pressure (SBP) significantly (p=0.026) in the trial conducted by Saltzman E et al. thus daily incorporation of 25-150g/d of oats or 2.5-6.0g of beta glucan is effective in the treatment of mildmoderate dyslipidemia.

# **Psyllium**

Psyllium seed husk also known as isabgol are the seeds of plant Plantago otava. They are indigestible by the human intestine, thus helps in holding large amounts of water and making the stools bulky and soft for easy defecation; therefore they are used in the treatment of constipation, Irritable bowel syndrome, diarrhea etc. As early as in 1998, the FDA already have approved a healthy claim on daily incorporation of 3-12grams of psyllium along with a low fat diet may reduce risk of heart disease<sup>57</sup>. The hypolipidemic effects of psyllium mucilage have been discussed by various epidemiological studies in table 7.

#### **To Summarize**

Various human clinical trials on psyllium<sup>58-</sup> suggest that this mucilage enriched food item is effective in lowering total cholesterol, LDL-c, triglycerides and improving the concentrations of good cholesterol HDL-c in the serum of normal and hypercholesterolemic subjects. Also the effects were more prominent in postmenopausal women as concluded by Ganji V et al. Psyllium was also effective in lowering Apo B100: ApoA1, TC: HDL. LDL: HDL and also increasing conc. of Apo A1; the apoprotein of HDL; thus reducing the primary and secondary ASCVD risk factors. Moreyra A, et al also concluded that 15g/d of psyllium intervention was as effective in lowering cholesterol as 20mg of simvastatin alone. Therefore, psyllium 5-15g/d is an effective adjunct therapy and may provide an alternative to drug therapy for mild- moderate hypercholesterolemic individuals.

# CONCLUSION

Dyslipidemia and ASCVD morbidity is increasing at an alarming rate nationwide and thus it becomes essential to prevent or treat dyslipidemia to regress the associated metabolic derangements. Fortunately today we can combat this situation initially through diet and physical activity before switching on to cholesterol lowering drug therapy. Merely by including the discussed hypolipidemic dietary agents in the required dosage as inferred from researches and recommendations i.e. oats 45-150g/d or beta glucan 2.5-6g/g; 20-25g of soy protein or 30-150mg of isoflavones; flaxseeds 20-50g of whole or partially defatted respectively; psyllium 5-15g per day along with a low fat NCEP followed diet and recommended physical exercise are significantly effective in improving serum lipid profile. The discussed food items may be effective in preventing, reversing or managing most prevalent chronic conditions as mentioned earlier, and hence may be claimed as to be "first line therapy" in dyslipidemia management. They can simply be incorporated in ones daily diet as they are less expensive, readily available, relatively have no side effects, practical and effective in recommended low dosages. They are proven to be more effective in mild-moderate hyperlipidemics like postmenopausal women who are hesitant to follow a life-long drug therapy for improving their mildly deranged lipid profile. This review attempts to not only enlist the effects, possible mechanism and bioactive components of the food item but also to quantify the same to have a desired effect on individual lipid profile.

# Limitations

While the majority of the published intervention studies suggest a lipid lowering effects of moderate doses of these food items administration, question remains about the

# REFERENCES

- National Commission on Macroeconomics and Health (NCMH) Background, Current Science, Vol. 97 (2005).
- 2. www.whoindia.org/en/index.
- National Cholesterol Education Program (NCEP) Expert Panel On Detection, Evaluation, and Treatment of High Blood Cholesterol In Adults (Adult Treatment Panel Iii) Final Report, Circulation 106, 31-43(2002).
- 4. AOAC Definition Revised By American Association of Cereal Chemists, 2000.
- Anderson JW, et al., Dietary Fiber Content of Selected Foods; American Journal of Clinical Nutrition vol.47, 440-7 (1988).
- C. Gopalan et al. Nutritive value of Indian foods, National Institute of Nutrition; 2011
- Brown Lisa, Bernard Rosner, Walter W Willett and Frank M Sacks. Cholesterol-Lowering Effects of Dietary Fiber: A Meta-Analysis. American Journal Of Clinical Nutrition, Vol. 69, No. 1, 30-42 (1999).
- 8. Anderson JW et al. Dietary Fiber: Hyperlipidemia, Hypertension and Coronary Artery Disease. Am J Gastroenterol, vol 81, issue 10, 907-919(1986).
- Schneeman Bo, Gallaher D. Effects of Dietary Fiber on Digestive Enzyme Activity and Bile Acids In The Small Intestine. Proc Soc Exp Biol Med; 180,409-414(1985).
- Jenkins DJ, Newton C, Leeds AR, Cummings JH (1975). Effect of Pectin, Guar Gum, and Wheat Fibre On Serum-Cholesterol. Lancet 1, 1116-1117.
- Hillman LC, Peters SG, Fisher CA, Pomare EW. The Effects of the Fiber Components Pectin, Cellulose and Lignin on Serum Cholesterol Levels. Am J Clin Nutr 42, 207-213(1985)
- 12. Schneeman BO. Dietary Fiber and Gastrointestinal Function .Nutrition Review

45, 129–32(1987)

premenopausal women).

 Blundell JE, Burley VJ. Satiation, Satiety and the Action of Fiber on Food Intake. Int J Obes (Suppl) 11, 9–25(1987).

consistency, efficacy, safety and duration of these

effects. Future studies will need to clarify the lipid

effects in more varied populations (men and

- Shearer G, Harris W, Pedersen T, Newman J. Detection of Omega-3 Oxylipins in Human Plasma and Response to Treatment with Omega-3 Acid Ethyl Esters. J Lipid Res. 51, 2074-2081(2009)
- 15. Holman R. The Slow Discovery of the Importance of Omega 3 Essential Fatty Acids in Human Health. J. Nutr. 128 (2 Suppl), 427s-433s(1998)
- Sanders T, Oakley F, Miller G, et al .Influence Of N-6 Versus N-3 Polyunsaturated Fatty Acids In Diets Low In Saturated Fatty Acids On Plasma Lipoproteins And Hemostatic Factors. Arterioscler Thromb Vasc Biol. 17, 3449–3460 (1997)
- Harris W, Connor W, Alam N, et al.. Reduction of Postprandial Triglyceridemia in Humans by Dietary N-3 Fatty Acids. J Lipid Res. 29, 1451–1460(1988)
- K.Prasad et al..Dietary Flax Seed in Prevention of Hypercholesterolemic Atherosclerosis. *Atherosclerosis* 132(1), 69-76(1997)
- Canadian Grain Commission. Nutritional profile of No. 1 Canada Western flaxseed and of yellow flaxseed samples. Canadian Grain Commission, Winnipeg, MB, 2001.
- Yildiz, Fatih. Phytoestrogens in Functional Foods. Taylor & Francis Ltd. Pp. 3–5, 210-211 (2005)
- Thompson L, Boucher B, Liu Z, Cotterchio M, Kreiger N. Phytoestrogen Content of Foods Consumed In Canada, Including Isoflavones, Lignans, and Coumestan. Nutrition and Cancer 54 (2), 184–201(2006)
- 22. Sacks F, Lichtenstein A, Van Horn L, Harris W, Kris-Etherton P, Winston M. Soy Protein,

Isoflavones, and Cardiovascular Health: An American Heart Association Science Advisory for Professionals from the Nutrition Committee. Circulation 113 (7): 1034– 44(2006)

- Turner J, Agatonovic S, Glass B. Molecular Aspects of Phytoestrogen Selective Binding at Estrogen Receptors. J Pharm Sci 96 (8), 1879–85(2007).
- Crouse J, Morgan Tm, Terry J, Ellis J, Vitolins M, Burke G. A Randomized Trial Comparing The Effect Of Casein With That Of Soy Protein Containing Varying Amounts Of Isoflavones On Plasma Concentrations Of Lipids And Lipoproteins. Arch Intern Med.159, 2070– 2076(1999)
- Wangen K, Duncan A, Xu X, Kurzer M. Soy Isoflavones Improve Plasma Lipids In Normocholesterolemic And Mildly Hypercholesterolemic Postmenopausal Women. Am J Clin Nutr. 73, 225–231(2001)
- 26. Goodman-Gruen D, Kritz-Silverstein D. Usual Dietary Isoflavone Intake Is Associated With Cardiovascular Disease Risk Factors In Postmenopausal Women. J Nutr. 131, 1202– 1206(2001)
- De Kleijn Mjj, Van Der Schouw Yt, Wilson Pwf, Grobbee De, Jacques Pf. Dietary Intake Of Phytoestrogens Is Associated With A Favorable Metabolic Cardiovascular Risk Profile In Postmenopausal Us Women: The Framingham Study. J Nutr. 132 no. 2 276-282 (2002)
- FAO/WHO Protein Quality Evaluation Report Of Joint FAO/WHO Expert Consultation, Food And Agriculture Organization Of The United Nations, FAO Food And Nutrition Paper No. 51, Rome. (1991)
- Schaafsma, G.. The Protein Digestibility-Corrected Amino Acid Score. Journal Of Nutrition 130, 1865s-1867s. (2000)
- Protein Quality-Report Of Joint FAO'/WHO Expert Consultation, Food And Agriculture Organisation, Rome, Fao Food And Nutrition Paper 51, 1991.
- Frank M. et al. Soy Protein, Isoflavones, And Cardiovascular Health. *Circulation* 113, 1034-1044. (2006)
- 32. Anderson J, Johnstone B, Cook-Newell M. Meta-Analysis of The Effects Of Soy Protein

Intake On Serum Lipids. N Engl J Med 333, 276–282. (1995)

- Carroll K. Review of Clinical Studies on Cholesterol-Lowering Response to Soy Protein. J Am Diet Assoc 91, 820-827. (1991)
- Washburn, Scott; Burke, Gregory L.; Morgan, Timothy; Anthony, Mary, Effect of Soy Protein Supplementation on Serum Lipoproteins, Blood Pressure, and Menopausal Symptoms in Perimenopausal Women. Menopause 6(1), 7-13. (1999)
- 35. <u>http://www.accessdata.fda.gov/scripts/cdrh/</u> <u>cfdocs/cfcfr/CFRSearch.cfm?fr=101.82;</u> Accessed on 25<sup>th</sup> July, 2013
- Pipe Elizabeth A et al..Soy Protein Reduces Serum LDL Cholesterol And The LDL Cholesterol:HDL Cholesterol And Apo lipoprotein B:Apolipoprotein A-I Ratios In Adults With Type 2 Diabetes . J. Nutr 139(9), 1700-06. (2009)
- Taku K et al.. Soy Isoflavones Lower Serum Total And LDL Cholesterol In Humans: A Meta-Analysis Of 11 Randomized Controlled Trials. American Journal Of Clinical Nutrition 85(4), 1148-1156. (2007)
- Clerici C et al.. Pasta Naturally Enriched With Isoflavone Aglycons From Soy Germ Reduces Serum Lipids And Improves Markers Of Cardiovascular Risk. J. Nutr. 137, 2270-2278. (2007)
- Desroches S et al. Soy Protein Favorably Affects Ldl Size Independently Of Isoflavones In Hypercholesterolemic Men And Women. J. Nutr. 134, 574-579. (2004).
- Wangen K, et al. Soy Isoflavones Improve Plasma Lipids In Normocholesterolemic And Mildly Hypercholesterolemic Postmenopausal Women. American Journal of Clinical Nutrition 73(2), 225-231. (2001).
- 41. Wofford MR et al. Effect of soy and milk protein supplementation on serum lipid levels: a randomized controlled trial. European Journal of Clinical Nutrition 66, 419– 425(2012)
- Flax Nutrition Profile Analyzed By the American Oil Chemists' Society (AOCS) Analysis by American Oil Chemist's Society (AOCS) Official Method Am 2-93 (2008.)
- 43. Pan A., Danxia Yu, Wendy Demark-Wahnefried, Oscar H Franco And Xu Lin.

Meta-Analysis of the Effects Of Flaxseed Interventions On Blood Lipids. *Am J Clin Nutr* **90**: 288-297. (2009)

- Patade A, et al Flaxseed Reduces Total and LDL Cholesterol Concentrations in Native American Postmenopausal Women. Journal of Women's Health 17(3): 355-366. (2008).
- Manda<sup>o</sup>escu S, Mocanu V, Dăscaliþa Am, Haliga R, Nestian I, Stitt Pa, Flaxseed Supplementation In Hyperlipidemic *Patients*. *Rev Med Chir Soc Med Nat Lasi*, **109**(3): 502-6. (2005).
- Jenkins DJ, et al.. Health Aspects of Partially Defatted Flaxseed, Including Effects on Serum Lipids, Oxidative Measures, And Ex Vivo Androgen and Progestin Activity: A Controlled Crossover Trial. *American Journal* of Clinical Nutrition 69(3): 395-402. (1999)
- Arjmandi, B.H *et al.* Whole Flaxseed Consumption Lowers Serum Ldl-Cholesterol And Lipoprotein (A) Concentrations In Postmenopausal Women. *Nutrition-Research* 18(7): 1203-1214. (1998).
- Fukumitsu S et al. Flaxseed lignan lowers blood cholesterol and decreases liver disease risk factors in moderately hypercholesterolemic men. Nutrition Research 30, 441–446(2010)
- FDA/CFSAN A Food Labeling Guide: <u>Appendix C Health Claims, April 2008</u>, Accessed on 25<sup>th</sup> July, 2013
- Villasmil R et al,. Oat-Derived [Beta]-Glucan Significantly Improves HDL-c And Diminishes LDL-c And Non-HDL Cholesterol In Overweight Individuals With Mild Hypercholesterolemia. American Journal of Therapeutics 14(2): 203-212. (2007)
- 51. Saltzman E. et al. An Oat-Containing Hypocaloric Diet Reduces Systolic Blood Pressure and Improves Lipid Profile Beyond Effects of Weight Loss In Men And Women. *Journal of Nutrition* 131, 1465-1470. (2001).
- 52. Romero A, et al.. Cookies Enriched With Psyllium Or Oats Lower Plasma LDL Cholesterol In Normal And Hypercholesterolemic Men From Northern Mexico. Journal Of The American College Of Nutrition **17**(6): 601-608. (1998)

- 53. Queenen K, et al..Concentrated Oats Beta Glucan, A Fermentable Fiber Lowrs (2007)Cholesterol in Hypercholesterolemic Adults in a Randomized Control *Trial. Nutrition Journal* 6: 525-524.
- 54. Robitaille J. et al,. Effect of an Oat Bran-Rich Supplement on the Metabolic Profile of Overweight Premenopausal Women. *Ann Nutr Metab* **49**: 141-148. (2005)
- 55. Maki et al. Whole-Grain Ready-to-Eat Oat Cereal, as Part of a Dietary Program for Weight Loss, Reduces Low-Density Lipoprotein Cholesterol in Adults with Overweight and Obesity More than a Dietary Program Including Low-Fiber Control Foods. J Am Diet Assoc.;110: 205-214 (2010)
- 56. Charlton K et al. Effects of 6 weeks consumption of B-glucan rich oat products on cholesterol levels in mildly hypercholesterolemic overweight adults. *British Journal of Nutrition*, **107**: 1037-1047(2012)
- 57. <u>http://www.accessdata.fda.gov/scripts/cdrh/</u> <u>cfdocs/cfcfr/CFRSearch.cfm?fr=101.81</u>, Accessed on 25<sup>th</sup> July, 2013
- Ganji Vijay and Jennifer Kuo. Serum Lipid Responses to Psyllium Fiber: Differences between Pre- and Post-Menopausal Hypercholesterolemic Women. *Nutrition Journal* 7(22): 7-22. (2008)
- Solà R, et al.. Effects of Soluble Fiber (Plantago Ovata Husk) On Plasma Lipids, Lipoproteins, and Apolipoproteins in Men with Ischemic Heart Disease American Journal of Clinical Nutrition 85(4): 1157-1163. (2007)
- Moreyra AE et al. Effect of Combining Psyllium Fiber with Simvastatin in Lowering Cholesterol. Archicves of Internal Medicine 165 (10): 1161-1166 (2005).
- 61. Anderson J W, et al. Long-Term Cholesterol-Lowering Effects of Psyllium as an Adjunct to Diet Therapy In The Treatment Of Hypercholesterolemia. *American Journal of Clinical Nutrition* **71**(6): 1433-1438
- Sartore G et al. The effects of psyllium on lipoproteins in type II diabetic patients. *European Journal of Clinical Nutrition* (2009)
  63: 1269–1271(2000).

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