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Pectin Composition, Food Applications, and Physiological Effects, Pectin Influences the Absorption, and Drug Delivery Applications Metabolism

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Abstract:

Dietary soluble fibre has gained consumer attention due to its supposed health benefits, which include lowering plasma cholesterol levels and slowing glucose absorption. Also getting a lot of attention is pectin, which is a major component of dietary fibre. The intrinsic features of pectin, variations in molecular weight across pectins, and the quantity of esterified methoxyl groups to galacturonic acid are some of the chemistry-related factors that draw attention to this soluble fibre. Research on pectin's physicochemical properties is mostly driven by the food industry's interest in the substance and its possible use in food products to enhance various qualities, such as water-holding capacity, gel consistency, and Scientists have speculated about the unique effects of pectin on lipid, viscosity. carbohydrate, mineral, and vitamin metabolism in the gastrointestinal system due to its physicochemical qualities. The unique chemical and biological characteristics of pectin have made it a hot topic in the field of biomedicine as of late. Pectin and other cellinstructive polymers are promising natural biomaterials for regenerative medicine. Furthermore, pectin-based composites and bioactive pectin both have enhanced properties that make them ideal for delivering active compounds. In vivo, pectin and pectin-based composites promote cell adhesion and proliferation, increase tissue remodelling, and function as interactive matrices or scaffolds. Tissue engineering and drug delivery systems benefit from pectin and pectin-based composites because of their immunoregulatory, antibacterial, anti-inflammatory, anti-tumor, and antioxidant actions, among other bioactive features. Nontoxicity, tunable mechanical qualities, biodegradability, and appropriate surface properties are critical elements of tissue engineering scaffolds that contain pectin or pectin-based conjugates or composites. Tissue engineering and medication distribution are two areas where pectic composites shine because to their adaptability in design and production.

Keywords: Food Applications, Metabolism, Pectin Composition, Drug Delivery

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Introduction

Many people in the food sector are curious about pectin, but there have been rumblings that the market could be glutted owing to rising manufacturing capacity and falling global demand. The global usage of pectin, however, remains between 18,000 and 19,000 metric tonnes. The public is very interested in learning which meals and food items contain a lot of pectin because there is a lot of evidence that it may lower plasma cholesterol levels and alter the glycaemic response. Apple pomace, sugar beetroot pulp and sunflower heads are significant sources of pectin, which varies from 15 to 25 g/100 g depending on the plant tissue. Climate, soils, and other variables, such as variety, extraction process, and fruit ripeness, can affect the quantity of pectin in fruits, including citrus fruits and other fruits [1, 2]. The quantities in juice are insufficient to give the promised health advantages. One of the most common sources is citrus fruits; however, pectin is made from citrus fruit peels rather than the actual fruits themselves. Technical issues have contributed to many previously reported estimates, which have overstated the pectin content of fruits and vegetables. Incorporating peel content, determining methods, using soluble pectin instead of total pectin, and, on rare occasions, using underripe fruits are all factors that contribute to these mistakes. As an example, lemons have been found to contain pectin levels ranging from 2.8 to 3.0 g/100 g. A more accurate measurement of 0.63 g/100 g of pectin was found in lemons that had been peeled. Peeled oranges contain about 6 grammes of pectin per kilogramme, according to Ross et al., who found 0.57 grammes of pectin per 100 grammes of oranges. Less than 1 mg/100 g is also the level seen in other fruits. In general, pectin is highly connected with citrus fruits, and the peels are the primary source of pectin [3-6]. As previously stated, the analysis was overestimated by 150% for lemons and by as much as 1000% for grapefruit due to the inclusion of peels. There have been suggestions for commercial utilisation of unconventional sources that are rich in pectin. Sunflower heads are one such source; to make a commercially viable product, two separate extractions are required. Insoluble protopectin is formed when pectin and cellulose are found in the central lamellae of plant cell walls. Pectin is produced when fruit ripens from protopectin. Additionally, pectic compounds are dissolved, leading to an increase in pectin that is soluble in water and oxalate. For cherries, the latter has been linked to less acid-soluble pectin. Also, although though polygalacturonase doesn't seem to be needed for pectin solubilisation, it nevertheless degrades cell wall pectin extensively and causes a drop in molecular weight. But when tomatoes had their polygalacturonase production halted using a gene-wrecking procedure, the dish wouldn't soften as it aged [7, 8]. Pectin methylesterase is an enzyme that plays a role in the initial stage of fruit ripening by catalysing the hydrolysis of methyl ester groups. Its activity results in a less methylated pectin, which is then used as a substrate by polygalacturonase. Pectin solubility and degree of connection may be affected by alterations to the galacturonan backbone's neutral side chains, according to reports. Although the structural composition of pectins is a linear chain of 1,4-linked α -d-galacturonic acid units, the presence of neutral sugars like l-rhamnose, dgalactose, and l-arabinose is found in nearly all pectins. The addition of methyl alcohol esterifies a few carboxyl groups. The pectins' characteristics are mostly dictated by their chain length and degree of esterification (DE). Pectins can be classified into two categories: high- and low-methoxyl. Between fifty and eighty percent of pectins are esterified, making them high-methoxylpectins. In the case where the esterification degree is less than 50%, a new class of polymers known as low-methoxylpectins is formed. The amount of esterification determines the kind of gel that pectins can produce. In contrast to low-methoxylpectins, which need a divalent cation like calcium to form gels, high-methoxylpectins can be prepared in acidic pH and with high sugar concentrations. Unlike other fibres like cellulose and lignin, pectins may retain calcium by absorbing it to their surface. This suggests a weak interaction with the divalent cation. During gel formation, dimerisation of polygalacturonate sequences is the principal mechanism for low-methoxyl pectin chain attachment [8, 9]. Inhibiting the competitive action of polygalacturonate blocks weakens the calcium pectate gel. Dimerisation of chains in a twofold conformation is also suggested by the current evidence as the process of crosslinking. When exposed to low pH and water activity, highmethoxylpectins gel, indicating that crosslinking occurs not at junction zones but rather in aggregates of varying extents. One polysaccharide that exhibits intriguing gel-forming properties is pectin, which can do it through a variety of processes depending on the degree of methylation.

The Role of Physicochemical Characteristics in Food

The gelling powers of pectin are highly advantageous in the jam industry. Confections, jams, jellies, and marmalades consume over half of the world's pectin. The most crucial quality that determines pectin's commercial value is its capacity to produce gels. There may be further uses for pectins as emulsifying agents when combined with proteins. Instead of relying solely on protein's emulsifying capabilities to create stable emulsions, creams and mayonnaise can also be made by forming complexes with polysaccharides. Pectin significantly enhanced the emulsifying capabilities of protein, according to Dalev and Simeonova. Furthermore, pectin does not add calories to food, thus using it as an emulsifier could be useful for low-calorie goods that are nonetheless nutritious [10, 11]. The usage of pectins as thickeners in food is expanding, and they also have some medicinal and biodegradable film manufacturing It seems that the product's consistency is affected by the amount of methylation of pectin. applications. Hydrophobycity and hydrogen bonding cause high-methoxylpectins to form an irreversible gel when heated. On the other hand, low-methoxyl pectin gels can be used for glazing, retorting, microwaving, sterilisation, and pasteurisation of food, as well as for jams and jellies made in bakeries, because their heat reversibility is an advantage [12]. Emulsion stability can be enhanced by pectin, in addition to its gel-forming, water-holding, and thickening capabilities. Although sugar beetroot pectin does not have good gelling properties, its physicochemical properties remain mostly unchanged when frozen and thawed, and it has a high water-holding capacity, it is still not used in food production capacity and low viscosity, thus it can have rather important applications in the formulation f certain products. Gels made with low-methoxylpectins have different strengths depending on the polysaccharide's molecular properties and the medium's calcium ion concentration. The strong water-holding capacity and viscosity of sunflower head residues, for instance, make them an excellent candidate for use in food applications. These residues are also rich in low-methoxyl pectin. Some properties of the gel's strength, including DE, can be impacted by the pectin's molecular weight (MW). According to Panchev et al., pectins with a DE of 57% to 58% produced the strongest gel. Low MW pectins can also produce powerful gels, since their MW declined as DE increased. Dairy products with pectins having MW more than 100,000 formed stronger gels, contradicting these findings [13, 14]. For strawberry jam, low-methoxyl pectin is preferable than high-methoxyl pectin because it improves consistency without significantly altering flavour. It is evident from these research that the MW and DE are important factors influencing the sensory and physical properties of pectins, depending on the product they are going to be used in. Concerning the acquisition of sufficient products with appropriate shelf life and product qualities guaranteeing high customer acceptability, the impact of pectin chemistry on these features becomes a significant concern.



Figure 1. Food substance physical property classification





Peptides and the Absorption of Minerals

Human investigations have failed to show any evidence of pectin-mineral interactions, despitethe fact that fibre sources containing pectin can mix with metal ions in vitro. For instance, the calcium, magnesium, phosphorus, and zinc balances have remained unchanged when dietary pectin levels range from 10 to 36 g/day. Despite no known impairment in iron absorption in normal persons, research with ileostomy patients have demonstrated that pectin may influence iron absorption. Just like these findings in humans, research on animals has likewise failed to show that pectin influences mineral absorption. Researchers have looked at how pectin influences mineral absorption in rats. Oral administration of 58Fe in conjunction with pectin was delivered to rats after they had been fed 2% pectin for 7 weeks [15, 16]. No effect on iron absorption or turnover was seen when dietary pectin was provided at the levels used in this investigation. In a separate investigation including iron supplementation in anaemic rats, researchers found no significant changes in iron availability between the control group and the rats given pectins of different DE or MW. Additionally, after incorporating a dosage of 58Fe into erythrocytes, the effects of these ectins on iron availability were studied in healthy, developing rats. When given 75% esterified pectin with an aMWof 89,000, rats absorbed 57% more iron than the control group (48%). Serum iron and transferrin saturation in haematocrit and liver were increased in the pectin-fed rats compared to the control group. Based on these results, it seems that pectin with low MW and high DE improves heme iron availability. To find out how various fibre sources, such as pectin, cellulose, oat bran, and wheat bran, impacted mineral absorption in rats, Galibois et al. conducted tests. Pectin seemed to improve the apparent absorption of calcium, iron, zinc, and magnesium compared to wheat or oat bran [17, 18]. Iron was better absorbed when 5 grammes of fibre was added rather than 10%. Given that pectin had negligible effects on mineral absorption when contrasted with wheat or oat bran consumption, the scientists reasoned that the source, not the quantity, of fiber influenced absorptive characteristics. Especially at the dosages used for human consumption, these investigations do not show that pectin will considerably alter mineral absorption.

Pectin and the Absorption of Vitamins

A small number of human and animal research have examined pectin's impact on the absorption vitamin E, carotenoids, and vitamin B12. People whose diets included items high incarotenoids were shown to have reduced plasma β -carotene concentrations when pectin was introduced to their diets. The observations presented here indicate that the inhibitory impact of pectin in the foods ingested might explain why people consuming meals rich in

carotenoids had lower plasma β -carotene levels than those taking β -carotene supplements [19]. These findings call for additional research to clarify pectin's function in carotenoids absorption. Vitamin E bioavailability was examined in rats given pectin dosages ranging from 0% to 8%. Vitamin E concentrations in the livers of rats given 8% and 6% pectin were lower than those of rats given 0% or 3% pectin after 8 weeks. The researchers found that vitamin E availability in rats was reduced by larger doses of pectin. Despite these findings, guinea pigs given pectin had higher levels of α-tocopherol in their low-density lipoprotein (LDL) compared to guinea pigs given a cellulose control diet. With the exception of the fibre type, the two diets were nutritionally equivalent. Also, thiobarbituric reactive substances (TBARS) production after copper incubation showed that guinea pigs given pectin had LDL particles that were less oxidatively sensitive [20]. Pectin has a protective effect in this study by reducing plasma α -tocopherol concentrations and boosting the resistance to oxidation of LDL. That pectin considerably reduced plasma LDL cholesterol contents in guinea pigs and that the LDL particles were cholesteryl ester deficient compared to those LDL produced from control animals could be an explanation for these contentious results. Since lipid (cholesteryl ester) oxidation is thought to happen before apolipoprotein B changes, most research examining the causes of LDL oxidation concentrate on this process; hence, a particle with lower concentrations of cholesteryl ester would be less susceptible to oxidation. Since the likelihood of LDL oxidation increases with the amount of time it stays in plasma, another element linked with its potential oxidation is the amount of time it stays in circulation. Due to an increase in LDL turnover and a decrease in the availability of LDL for oxidation, pectin consumption enhanced LDL apo B/E receptors in guinea pigs. Pectin has been found to have a negative impact on vitamin B12 status in rats who are deficient in this nutrient. 57[Co] vitamin B12 had a 58-day biological life in the fiber-free group and a 38-day life in the pectin-fed rats. Methylmalonic acid, a byproduct of vitamin B12 breakdown, was also shown to be higher in urine after pectin The same authors later found that vitamin B12-deficient rats had higher methylmalonic acid consumption. concentrations than predicted from propionate synthesis from pectin, indicating that this fibre may affect vitamin absorption.

Controlling Blood Sugar Using Pectin

Postprandial glucose levels increase in response to an increase in blood insulin following a carbohydrate-containing meal. People with diabetes must pay close attention to these changes in glucose and insulin levels. Evidence suggests that pectin may help insulin-dependent and type II diabetes individuals control these alterations. A number of clinical trials have shown decreased plasma glucose concentrations. While some research has demonstrated that pectin inhibits the postprandial increase in insulin and glucose concentrations, other investigations have found the opposite to be true. Pectin was one of several fibre sources that Jenkins et al. evaluated on a sample of healthy volunteers. Adding each fibre to a glucose tolerance test resulted in lower blood glucose and insulin concentrations at various times. Neither cholestyramine nor bran had this effect. According to the scientists, the success in lowering plasma glucose levels was found to be closely correlated with the fiber's viscosity [21, 22]. Other research has demonstrated that healthy volunteers' plasma glucose concentrations can be decreased by adding pectin to test meals or glucose solutions. Pectin reduced the blood glucose response in healthy volunteers but had no impact on patients who had undergone total gastric bypass. Consistent with the previous findings, a research conducted by Schwartz et al. on type II diabetes patients found that sustained pectin administration increased stomach emptying half-time by 43% and improved glucose tolerance. All of these investigations point to the possibility that pectin's lower absorption and enhanced glycaemic response in both healthy people and diabetic patients are due, in part, to its increased viscosity in the gastrointestinal lumen. After being fed 5% citrus fibre, rats were studied to determine the effects of pectin on mucin, a high molecular weight glycoprotein that is responsible for the gel-like texture of intestinal mucus. An antirat mucine polyclonal antibody showed much higher reactivity in luminal samples from the stomach and intestines of the citrus fiber-fed group. More fast transit time and delayed or reduced nutritional absorption are among the known side effects of pectin feeding, and the data imply that these could be attributable to increases of gastrointestinal mucus. In addition, after 5 weeks of a pectin-supplemented diet, rats showed changes in their intestinal loops that were linked to lower glucose absorption. However, the investigators couldn't find any changes in the intestinal wall that could explain this reduced absorption. Additionally, after a three-day suspension of pectin supplementation, patients with non-insulin-dependent diabetes have demonstrated adaptive alterations in reaction to pectin feeding. In conclusion, these animal and human studies suggest that pectin may reduce the postprandial increase in blood glucose through two

distinct mechanisms: (a) changing the viscosity, which delays gastric emptying and intestinal absorption, and (b) causing changes in the intestinal barrier layer.

The Metabolism and Absorption of Pectin

One of the many health benefits of the polyphenols included in fruits and vegetables is their ability to ward against cardiovascular disease. Polyphenolic compounds' antioxidant capabilities, which include the ability to scavenge free radicals, chelate metals, and activate enzymes, are thought to be responsible for their biological activity. Polyphenol bioavailability varies greatly depending on factors such polyphenol structure and conjugation, dietary matrix components, and interactions with the gastrointestinal system. A compound's in vivo action is dependent on its metabolism, the kind and amount of its absorption and elimination, and the activity of other compounds in circulation. Several fruits and vegetables contain flavonoids, a diverse group of polyphenols. When flavonoids are broken down in the small intestine and liver, the unabsorbed ones are converted into phenolic acids through a process called microbiota-mediated ring-fission. These acids are then absorbed and eliminated from the body in the urine. Though polyphenols' health advantages have long been linked to the food-based parent substances, it is possible that the presence of their metabolites in tissues and blood is what actually imparts these biological features [23, 24]. Both directly and indirectly, via the gut bacteria, polyphenols can affect cellular processes. Around 5-10% of all polyphenols make it into the small intestine for absorption; the rest make it to the large intestine, where they are either broken down by resident bacteria or eliminated in the stool. The physicochemical characteristics of dietary fibre improve gut health and general wellness, making it a vital nutrient for optimal health. Our prior research in rats shown that the combined effects of fermentable fibre and blackcurrant were more beneficial to their health than either supplement alone. A balanced diet includes both macro and micronutrients, creating a complex dietary matrix that may influence the intestinal bioavailability of polyphenols. Dietary fibre may regulate the availability of polyphenols in the upper and lower gastrointestinal tracts, according to previous research.

Pectin: A Review of Its Biological Characteristics and Biomedical Uses

The natural polymer pectin has recently attracted a lot of interest from researchers because of its many potential pharmacological and therapeutic uses, as well as its cheaper price tag. Pectin and pectin-based composites are currently the subject of intense research due to their potential therapeutic effects, low toxicity, and use in food, healthcare, and cosmetics. Advancements in manufacturing, purification, and characterisation techniques, as well as new and improved in vitro and in vivo testing methods for influencing immunity, have also played a significant role in this ongoing investigation. Many Food and Agriculture Organisations have given their stamp of approval to commercially available pectin for certain uses, and it meets all of the necessary standards. Pectin is extracted commercially from the byproducts of fruit and vegetable processing, specifically from sugar beets and citrus fruits, after the juice and pulp have been extracted [25, 26]. For plants to grow, pectin-a component of the cell wall-is necessary. Drug delivery and tissue engineering are two areas where pectin has found efficient applications in the form of gel beads or microspheres, 3D scaffolds, and membranes. The three main pectic polysaccharides found in primary cell walls are homogalacturonan, rhamnogalacturonan-I, and rhamnogalacturonan-II. The structural complexity and heterogeneity of the pectin domains are caused by pectin-modifying enzymes and endomembrane system biosynthesis. Homogalacturonan can be modulated by the enzyme pectin methyl esterase. In contrast to rhamnogalacturonan-I, which displays a highly unstable pectin matrix, rhamnogalacturonan-II contains very different functionally controlled polymers. The main cell walls of terrestrial plants include pectin, a structurally related polymer rich in D-galacturonic acid that serves multiple purposes. The saccharide polymers-carrying remains of 1-2 connected alpha-L-rhamnopyranosyl are interchangeable with the covalently linked 1-4-alpha-D-galacturonic units. Most of the galacturonic residues in pectin exist as methyl esters or salts. Determining pectin's exact chemical structure is a challenging task. Their chemical arrangement varies depending on the source and conditions they extract in relation to geography and other surrounding factors. Fruits and vegetables like sugar beets, citrus peel, and apple pomace are the commercial sources of pectin. The polysaccharide contributes to the mechanical resistance, stiffness, and intercellular adhesion of plant cell walls. This is essential for the survival of plants in areas where temperature, pollution, and other environmental stresses are significant threats. Pectin offers a plethora of target sites for chemical alterations due to its multifunctional component. Alpha-1, 2-L-rhamnose units and alpha-1, 4-linked Dgalacturonic acid units alternate throughout the pectin polymer's central structure. The pectin system controls the polysaccharide's effect on cytokine production, proving that the polysaccharide's elemental properties are associated with its capacity to affect cellular environmental factors. Pectin polymers have a wide range of uses due to the structural variety found in them from various plant sources. Although the structures of pectin from different plants are generally similar, they do vary depending on the species and the stage of the plant's physiological development. The functioning of the polymer is determined by the chemical composition of pectin, which includes structural properties, the proportion of galacturonic acid, the presence of methyl groups, and the grade of acetylation. Biomedical and tissue engineering/drug delivery pectin uses are affected by immunological reactivity. Immunomodulators are crucial in the treatment of pathological disorders because they control the body's unique immune response to invaders and antigens introduced by foreign or transplanted cells [24, 25]. The purpose of utilizing immunomodulators such as pectin is not to eliminate the immune response but to regulate the reactivity and further the efficiency of the applications that require the modulation of the immune system. Past studies have reported that pectin can weaken inflammatory reactivity by stimulating anti-inflammatory cytokines and decreasing the assembly of proinflammatory cytokines. Due to its structural complexity and diversity, pectin has many applications. Pectin consists of many active functional groups of polysaccharides, enabling them to have much more excellent modification properties than other biopolymers. Pectin is a hydrophilic natural polymer that can absorb or retain much water and exhibit swelling properties. Hydrogels and composite materials can be formed by crosslinking and other techniques, and the matrix structure can be incorporated with various bioactive compounds. To add to their suitability as a biodegradable and biocompatible delivery system for bioactive compounds, pectin-based smart composites possess physical-sensitive (light, temperature, electricity), chemical-sensitive (pH, redox, glucose), and biologically-sensitive (enzymes) properties. Due to its broad availability, pectin has become a prominent branch of the research and development of nature-based biomedical and healthcare areas.



Figure 3. The biomedical uses of pectin and composites made of pectin are numerous.

Pectin for Use in Medicine Delivery

Due to their superior features, such as biocompatibility, nontoxicity, flexibility in production, and functionalization, natural polymer pectin hydrogels have earned substantial consideration in drug delivery applications. In the drug delivery system, the degradation rate of the hydrogel carrier is significant for delivering the active material to the target site. Using pectin within the medication delivery system has largely been researched since pectin hydrogels can release medicines. Generally, researchers in the drug delivery system desire the drug to be securely and efficiently immobilized or covalently bonded to a biomaterial vehicle such as pectin. Industrially, when integrated into a medication component, pectin is generally utilized to treat radioactive isotopes and heavy metal poisoning [26, 27]. Concerning heavy metals, pectin can operate as a chelating agent by eliminating or blocking the interactions of dangerous heavy metals within the human body, such as iron, copper, and mercury. When pectin polysaccharides interact with metal ions, esterification, and chelation occur according to the number of non-methyl-esterified galacturonosyl residues. Pectin molecules that are not esterified can form gels when surrounded by bivalent cations.



Figure 4. The medicine is delivered to the target tissue in vivo through the use of pectin or microspheres made of pectin, which release the drug at a predetermined rate.

Ionic crosslinks between galacturonan chains with six or more adjacent residues cause pectin molecules to broaden their metal binding and lower the degree of methyl esterification. The amount of galacturonic acid that undergoes esterification when reacting with methanol is known as the degree of esterification. Pectin is often grouped into two types: methoxyl pectin and high-methoxyl pectin. Ionic crosslinking between the homogalacturonan chains gives lowmethoxyl pectin the ability to form a gel when it is surrounded by calcium. Among the few known gelation mechanisms, the egg-box mechanism stands out. Between each homogalacturonan chain that is generated in the eggbox process, there are six or more adjacent non-esterified galacturonic residues in the calcium-crosslinked junction zones. An absorbent polymer network is created as a consequence of this interaction. The physical characteristics and rate of gelation of pectin polysaccharides are both affected by the extent to which the galacturonosyl residues have been methyl esterified. A number of established applications in fields as diverse as medicine, physics, chemistry, biotechnology, biochemistry, and cryobiology make use of pectin polysaccharides, one of the key reasons for this being their gel-forming capability. Pectin is a nutritive food ingredient, and polysaccharides are a class of medications used to treat gastrointestinal disorders [26, 27]. Digestible pectin is not an unmodified variety. The digestive system's motility and peristalsis are stimulated by pectin in humans. When muscles contract, a process called peristalsis Because of its gel-forming characteristics, pectin can enhance the absorption of both food and happens. physiologically stimulated substances, and it can also cleanse the villi of the small intestine. Both hydrogel-based drug delivery systems and medicines tailored to the colon have made use of pectin as a transport mechanism. In their most basic form, hydrogels are three-dimensional networks made of crosslinked, hydrophilic polymer chains. As a drug delivery vehicle, pectin in the hydrogel-based drug delivery system can release the targeted medication into the body at a specified pace and location [28, 29]. Through the use of polymers, certain medications can be directed to the colon rather than the upper intestines, a process known as colon-specific delivery. Additionally, the polymer can regulate the release of medications to meet certain target rates. Composite materials utilizing pectin have found new uses in fields like tissue engineering, thanks to the extensive interactions it forms with a wide variety of biopolymers.

Blood Concentrations of LDL Cholesterol and Pectin

The inherent properties of soluble fiber, which reduce the risk of coronary heart disease by lowering plasma LDL cholesterol and unbalancing nutrient absorption in the intestinal lumen, likely explain why increasing dietary consumption of this fiber is considered a protective habit against the risk of ischemic heart disease. A number of epidemiological studies have shown that dietary fiber may provide some protection against coronary heart disease, which sparked the current interest in this relationship between diet and cardiovascular disease. There is strong evidence that a diet high in fiber foods such fruits, vegetables, and grains reduces the risk of myocardial infarction. A higher fiber diet is an essential component in reducing the risk of coronary heart disease, therefore it stands to reason that recommendations should be made to that effect. Reduced hepatic cholesterol concentrations are a result of fiber

activity in the intestinal lumen (primary processes), which in turn cause significant changes in lipoprotein synthesis, intravascular processing, and catabolism (secondary mechanisms). Soluble fiber's effect varies with fiber type and dietary cholesterol intake. Here we will provide a concise overview of pectin's effects on human plasma lipid levels and the processes of decreasing plasma LDL cholesterol in animal models. A review of the literature found that pectin consumption lowers plasma LDL cholesterol levels but has no discernible influence on HDL cholesterol or triglyceride levels. The ways in which pectin works are a matter of some debate [30]. The majority of pectin digestion occurs in the colon, rather than the stomach or small intestine, according to feeding experiments with ileostomy patients. This data, along with the fact that galacturonic acid is not present in stool samples, lend credence to the idea that pectin is fermented by bacteria in the colon to produce short-chain fatty acids. These acids can then be used by the bacteria in the colon or absorbed by the mucosa lining the intestines. Propionic acid is the primary shortchain fatty acid generated, and these results imply that it may affect cholesterol production. Studies in rats, hamsters, and guinea pigs that used pectin or other forms of soluble fiber did not support this theory because the results showed an increase in hepatic or total body cholesterol synthesis following soluble fiber consumption, rather than a decrease. According to Levrat et al., rats whose diets included pectin had significantly higher levels of volatile fatty acid in their cecums. Consistent with the aforementioned studies, this one also found a marked upregulation of hepatic HMG-CoA reductase activity, the enzyme responsible for controlling cholesterol synthesis. This lends credence to the idea that the conversion of pectin to volatile fatty acids during colon fermentation does not lead to a reduction in cholesterol biosynthesis. It is possible that pectin lowers plasma cholesterol levels through additional pathways involving the intestinal lumen. Disruption of micelle production could be one mechanism. If the emulsion of mixed micelles, which is the sole way for dietary lipids to be absorbed, breaks down, then the lipid components will also be inaccessible. Reduced lipid absorption may occur if pectin destabilizes micelles, trapping or disintegrating them. The scientists who discovered that aluminum pectate significantly reduced rat plasma cholesterol speculated that this was due to the micelle's negative charges causing a complex to form with the positively charged aluminum. As mentioned before, pectin's viscosity could be involved in an additional method. The apparent thickness of the unstirred water layer increases with increasing viscosity, which in turn increases the resistance to diffusion and absorption. Both people and rats given 0-15 g/L of pectin showed a reduction in the absorption of glucose and fatty acids. As the concentration of pectin in the solution was raised, the viscosity also rose. Similar to bile acid-binding resins, pectin can enhance bile acid excretion, albeit to a lesser extent. Instead of pectin directly attaching to bile acids, this process might be associated with the physicochemical features of pectin that prevent bile acid circulation via the enterohepatic circulation. However, pectin's hypocholesterolemic effects have not been confirmed in all clinical trials. When healthy, normolipidemic volunteers consumed 12 grams of pectin daily, Hillman et al. did not observe a statistically significant decrease in plasma cholesterol. The authors speculated that the lack of impact might be due to the fact that the participants used were either normal-cholesterolemic or not eating a diet heavy in cholesterol. After four weeks of eating citrus fiber, Wisker et al. found that women's plasma cholesterol levels dropped by 11%. But since HDL cholesterol was the only kind of plasma cholesterol that was reduced, the total-to-HDL cholesterol ratio did not improve.

Benefits of Pectin

Many different health concerns motivate people to take MCP. However, animal experiments provide the bulk of our knowledge regarding pectin.

Pectin for cholesterol

Pectin, along with other soluble fibers like oatmeal and psyllium husks, may aid in lowering LDL ("bad") cholesterol, according to some research. However, it has a negligible impact. In certain cases, soluble fibers like pectin may assist reduce cholesterol levels; however, they are rarely effective when used alone.

Pectin for diarrhea

Some early antidiarrheal medications relied on pectin as an active component to manage diarrhea. When it comes to treating infants and toddlers, there is some evidence that it works. However, in 2003, the FDA determined that there was insufficient data to justify this use. It outlawed pectin in over-the-counter diarrhea remedies the year after that.

Pectin for cancer

One possible use for pectin is in cancer treatment. A tiny research found that MCP appeared to reduce the progression of prostate cancer in men who had failed with normal treatment. In a study that followed men for a longer period of time, researchers found that MCP reduced PSA levels in men who had a specific kind of prostate cancer. Cancers of the breast and prostate may be able to halt their metastasis (the spread of cancer cells to other parts of the body) if MCP is used. Research has revealed that a different type of pectin can inhibit the growth of pancreatic cancer cells by blocking important processes.

Pectin for metal toxicity

A number of heavy metal toxins, including lead, mercury, arsenic, and others, have been treated with pectin. There are others who think MCP can aid in the elimination of these harmful compounds from the body. However, there is a lack of impartial research to back up these assertions.

There is no known ailment for which the appropriate dose of MCP has been determined. The active chemicals in MCP products can vary in quality from one manufacturer to another, as is typical with supplements in general. The Food and Drug Administration has not yet green-lit MCP for the treatment of any illness.

Side Effects of Pectin

Taking MCP is connected with little negative effects. But that doesn't imply there isn't any danger.

- Mild gastrointestinal cramps and diarrhea have been reported by some individuals when using MCP.
- Stay away from MCP if you have a citrus fruit allergy.

You shouldn't take MCP without a doctor's supervision because it might counteract the effects of some cancer medicines.

The absorption of the vital vitamin beta-carotene can be hindered by pectin. Pectin can also affect the absorption of some medications, such as: • Digoxin, which is used to treat cardiac conditions; • Lovastatin, which is used to control cholesterol levels in the body.

Drugs belonging to the tetracycline family

Dietary supplements are not subject to the same level of regulation by the FDA as meals and medications. Ensuring safety and correct labeling falls on the manufacturer. Before taking pectin or any other dietary supplement, it is important to discuss the dangers with your doctor.

Pectin vs. Gelatin

There are a few key distinctions between pectin and gelatin, two thickeners that find common use in food and medicine. Where it all began. Plants are the sources of pectin, while animal collagen is the source of gelatin. Collagen is present in skin, bones, and other tissues. When cooked with sugar and acid, pectin produces a thick and sticky gel. That's why it works wonderfully with jam. Commercial yogurt smoothies, protein drinks, and syrups can all benefit from pectin's thickening properties. The texture of gelatin becomes smoother and creamier after it is boiled and chilled. Jell-O relies on gelatin, which is also an ingredient in marshmallows, ice cream, mousse, and cheese.

CONCLUSION

Although there may be some variation linked with dosages employed, the inclusion of hyperlipidemic patients, or the animal model examined, the majority of human and animal research have demonstrated that pectin decreases plasma cholesterol concentrations. It appears that pectin's water-holding capacity, viscosity, and possible gel formation are related to its action, according to the evidence that is now available. Delays in stomach emptying, reduced mobility through the ileum (the site of lipid absorption), and interference with micelle production are all outcomes of these characteristics. Pectin improves bile acid excretion but does not bind to bile acids directly; this disrupts the bile acid enterohepatic circulation. When pectin reaches the colon, the bacteria there have already broken it down into short-chain fatty acids. It is currently unclear how these volatile fatty acids contribute to fiber's hypocholesterolemic effect. The idea that volatile fatty acids reduce cholesterol production is refuted by investigations done in various animal models showing that pectin upregulates hepatic cholesterol biosynthesis in response to the depletion of liver pools of

cholesterol. Therefore, it is reasonable to assume that pectin's primary action—a reduction in plasma cholesterol occurs in the small intestine and stomach, prior to its degradation and fermentation in the large intestine. Hepatic cholesterol concentrations are reduced because pectin either creates a physical barrier that delays cholesterol absorption or disrupts micelle formation. Alternatively, it promotes mobilization of hepatic cholesterol by enhancing bile acid synthesis and interrupts the enterohepatic circulation of bile acids. Hepatic apo B/E receptors are activated by pectin, which leads to a shift in cholesterol homeostasis and the subsequent removal of cholesterol from plasma. Thus, pectin is a soluble fiber that, in most cases, lowers plasma LDL cholesterol levels. It may also directly affect the reduction of arteriosclerosis, as demonstrated in animal experiments. Because of its many positive effects, pectin consumption may improve general health and lower the risk of developing degenerative disorders like diabetes and coronary heart disease. The biological advantages of pectin and pectin-based composites include their lack of toxicity, compatibility with living organisms, ability to break down naturally, and potential to fight against cancer, bacteria, and tumors. Drug delivery and tissue engineering are two areas where these materials could be particularly useful due to their high chemical reactivity and emulsification capabilities. Research into the potential medicinal uses of functionalized scaffolds made of natural macromolecular materials is underway, with applications in the fields of skin, biological valves, bone tissue, and injectable scaffolds. Nevertheless, additional research is necessary to delve into the functions of pectin's bioactivity in living organisms. Clinical trials and research into pectin's anticancer protective mechanisms are lacking, which limits the substance's potential medical and pharmaceutical uses. Degradation kinetics in both vitro and vivo, data on digested products, and action mechanisms could shed light on how pectin can be technologically advanced to increase its clinical use in tissue engineering and biomedicine.

REFERENCES

- 1. Li, Y.O.; Komarek, A.R. Dietary fibre basics: Health, nutrition, analysis, and applications. Food Qual. Saf. 2017, 1, 47–59.
- 2. Paturi, G.; Butts, C.A.; Monro, J.A.; Hedderley, D. Effects of blackcurrant and dietary fibers on large intestinal health biomarkers in rats. Plant Foods Hum. Nutr. 2018, 73, 54–60.
- 3. Bohn, T. Dietary factors affecting polyphenol bioavailability. Nutr. Rev. 2014, 72, 429–452
- 4. Aguilera, J.M. The food matrix: Implications in processing, nutrition and health. Crit. Rev. Food Sci. Nutr. 2019, 59, 3612–3629.
- 5. Jakobek, L.; Mati'c, P. Non-covalent dietary fiber—Polyphenol interactions and their influence on polyphenol bioaccessibility. Trends Food Sci. Technol. 2019, 83, 235–247.
- Mullen, W.; Edwards, C.A.; Crozier, A. Absorption, excretion and metabolite profiling of methyl-, glucuronyl-, glucosyl- and sulpho-conjugates of quercetin in human plasma and urine after ingestion of onions. Br. J. Nutr. 2006, 96, 107–116.
- 7. Gee, J.M.; Wroblewska, M.A.; Bennett, R.N.; Mellon, F.A.; Johnson, I.T. Absorption and twenty-four-hour metabolism time-course of quercetin-3-O-glucoside in rats, in vivo. J. Sci. Food Agric. 2004, 84, 1341–1348.
- 8. Valkonen, M.; Kuusi, T. Spectrophotometric assay for total peroxyl radical-trapping antioxidant potential in human serum. J. Lipid Res. 1997, 38, 823–833.
- 9. Pires, M.A.; Pastrana, L.M.; Fuciños, P.; Abreu, C.S.; Oliveira, S.M. Sensorial perception of astringency: Oral mechanisms and current analysis methods. Foods 2020, 9, 1124.
- 10. Van Hul, M.; Cani, P.D. Targeting carbohydrates and polyphenols for a healthy microbiome and healthy weight. Curr. Nutr. Rep. 2019, 8, 307–316
- 11. Liu, Y.; Weng, P.; Liu, Y.; Wu, Z.; Wang, L.; Liu, L. Citrus pectin research advances: Derived as a biomaterial in the construction and applications of micro/nano-delivery systems. Food Hydrocoll. 2022, 133, 107910.
- 12. Coimbra, P.; Ferreira, P.; de Sousa, H.C.; Batista, P.; Rodrigues, M.A.; Correia, I.J.; Gil, M.H. Preparation and chemical and biological characterization of a pectin/chitosan polyelectrolyte complex scaffold for possible bone tissue engineering applications. Int. J. Biol. Macromol. 2011, 48, 112–118.

- 13. Neufeld, L.; Bianco-Peled, H. Pectin-chitosan physical hydrogels as potential drug delivery vehicles. Int. J. Biol. Macromol. 2017, 101, 852–861.
- 14. Pereira, R.F.; Barrias, C.C.; Bártolo, P.J.; Granja, P.L. Cell-instructive pectin hydrogels crosslinked via thiol-norbornene photo-click chemistry for skin tissue engineering. Acta Biomater. 2018, 66, 282–293.
- 15. Khotimchenko, Y.; Khozhaenko, E.; Kovalev, V.; Khotimchenko, M. Cerium binding activity of pectins isolated from the seagrasses Zostera marina and Phyllospadixiwatensis. Mar. Drugs 2012, 10, 834–848.
- 16. Pérez, S.; Mazeau, K.; Hervé du Penhoat, C. The three-dimensional structures of the pectic polysaccharides. Plant Physiol. Biochem. 2000, 38, 37–55.
- 17. Marisol, O.-V.; Emmanuel, A.-H.; Irasema, V.-A.; Miguel ÁngelMarisol, M.-T. Plant Cell Wall Polymers: Function, Structure and Biological Activity of Their Derivatives. In Polymerization; Ailton De Souza, G., Ed.; IntechOpen: Rijeka, Croatia, 2012; p. 4.
- 18. Cuijpers, V.M.; Walboomers, X.F.; Jansen, J.A. Scanning electron microscopy stereoimaging for three-dimensional visualization and analysis of cells in tissue-engineered constructs: Technical note. Tissue Eng. Part. C Methods 2011, 17, 663–668.
- 19. O'Brien, F.J. Biomaterials & scaffolds for tissue engineering. Mater. Today 2011, 14, 88-95.
- 20. Kumar, P.T.; Ramya, C.; Jayakumar, R.; Nair, S.K.; Lakshmanan, V.K. Drug delivery and tissue engineering applications of biocompatible pectin–chitin/nano CaCO₃ composite scaffolds. Colloids Surf. B Biointerfaces 2013, 106, 109–116.
- 21. Salman, H.; Bergman, M.; Djaldetti, M.; Orlin, J.; Bessler, H. Citrus pectin affects cytokine production by human peripheral blood mononuclear cells. Biomed. Pharmacother. 2008, 62, 579–582.
- Popov, S.V.; Markov, P.A.; Popova, G.Y.; Nikitina, I.R.; Efimova, L.; Ovodov, Y.S. Antiinflammatory activity of low and high methoxylated citrus pectins. Biomed. Prev. Nutr. 2013, 3, 59– 63.
- 23. Boehler, R.M.; Graham, J.G.; Shea, L.D. Tissue engineering tools for modulation of the immune response. Biotechniques 2011, 51, 239–240, 242, 244.
- 24. Daguet, D.; Pinheiro, I.; Verhelst, A.; Possemiers, S.; Marzorati, M. Arabinogalactan and fructooligosaccharides improve the gut barrier function in distinct areas of the colon in the Simulator of the Human Intestinal Microbial Ecosystem. J. Funct. Foods 2016, 20, 369–379.
- 25. Vogt, L.M.; Sahasrabudhe, N.M.; Ramasamy, U.; Meyer, D.; Pullens, G.; Faas, M.M.; Venema, K.; Schols, H.A.; De Vos, P. The impact of lemon pectin characteristics on TLR activation and T84 intestinal epithelial cell barrier function. J. Funct. Foods 2016, 22, 398–407.
- 26. Ho, G.T.; Zou, Y.F.; Aslaksen, T.H.; Wangensteen, H.; Barsett, H. Structural characterization of bioactive pectic polysaccharides from elderflowers (Sambuciflos). Carbohydr. Polym. 2016, 135, 128–137.
- 27. Kapoor, S.; Dharmesh, S.M. Pectic Oligosaccharide from tomato exhibiting anticancer potential on a gastric cancer cell line: Structure-function relationship. Carbohydr. Polym. 2017, 160, 52–61.

- 28. Liu, Z.; Dang, J.; Wang, Q.; Yu, M.; Jiang, L.; Mei, L.; Shao, Y.; Tao, Y. Optimization of polysaccharides from Lyciumruthenicum fruit using RSM and its antioxidant activity. Int. J. Biol. Macromol. 2013, 61, 127–134.
- 29. Peng, Q.; Xu, Q.; Yin, H.; Huang, L.; Du, Y. Characterization of an immunologically active pectin from the fruits of Lyciumruthenicum. Int. J. Biol. Macromol. 2014, 64, 69–75.
- Leivas, C.L.; Nascimento, L.F.; Barros, W.M.; Santos, A.R.; Iacomini, M.; Cordeiro, L.M. Substituted galacturonan from starfruit: Chemical structure and antinociceptive and antiinflammatory effects. Int. J. Biol. Macromol. 2016, 84, 295–300.
- 31. Luan, F.; Peng, L.; Lei, Z.; Jia, X.; Zou, J.; Yang, Y.; He, X.; Zeng, N. Traditional Uses, Phytochemical Constituents and Pharmacological Properties of Averrhoa carambola L.: A Review. Front. Pharmacol. 2021, 12, 699899.
- 32. Oueslati, S.; Ksouri, R.; Falleh, H.; Pichette, A.; Abdelly, C.; Legault, J. Phenolic content, antioxidant, anti-inflammatory and anticancer activities of the edible halophyte SuaedafruticosaForssk. Food Chem. 2012, 132, 943–947.
- 33. Mzoughi, Z.; Abdelhamid, A.; Rihouey, C.; Le Cerf, D.; Bouraoui, A.; Majdoub, H. Optimized extraction of pectin-like polysaccharide from Suaedafruticosa leaves: Characterization, antioxidant, anti-inflammatory and analgesic activities. Carbohydr. Polym. 2018, 185, 127–137.
- 34. Sherry, C.L.; Kim, S.S.; Dilger, R.N.; Bauer, L.L.; Moon, M.L.; Tapping, R.I.; Fahey, G.C., Jr.; Tappenden, K.A.; Freund, G.G. Sickness behavior induced by endotoxin can be mitigated by the dietary soluble fiber, pectin, through up-regulation of IL-4 and Th2 polarization. Brain Behav. Immun. 2010, 24, 631–640.
- 35. Do Nascimento, G.E.; Winnischofer, S.M.B.; Ramirez, M.I.; Iacomini, M.; Cordeiro, L.M.C. The influence of sweet pepper pectin structural characteristics on cytokine secretion by THP-1 macrophages. Food Res. Int. 2017, 102, 588–594.
- 36. Pedrosa, L.F.; Raz, A.; Fabi, J.P. The Complex Biological Effects of Pectin: Galectin-3 Targeting as Potential Human Health Improvement? Biomolecules 2022, 12, 289.
- 37. Zhang, W.; Zhao, X.J.; Jiang, Y.; Zhou, Z. Citrus pectin derived silver nanoparticles and their antibacterial activity. Inorg. Nano-Met. Chem. 2017, 47, 15–20.
- 38. Gupta, V.K.; Pathania, D.; Asif, M.; Sharma, G. Liquid phase synthesis of pectin–cadmium sulfide nanocomposite and its photocatalytic and antibacterial activity. J. Mol. Liq. 2014, 196, 107–112.
- 39. Pathania, D.; Sharma, G.; Thakur, R. Pectin @ zirconium (IV) silicophosphate nanocomposite ion exchanger: Photo catalysis, heavy metal separation and antibacterial activity. Chem. Eng. J. 2015, 267, 235–244.
- 40. Hassan, E.A.; Abou Elseoud, W.S.; Abo-Elfadl, M.T.; Hassan, M.L. New pectin derivatives with antimicrobial and emulsification properties via complexation with metal-terpyridines. Carbohydr. Polym. 2021, 268, 118230.
- 41. Supreetha, R.; Bindya, S.; Deepika, P.; Vinusha, H.; Hema, B. Characterization and biological activities of synthesized citrus pectin-MgO nanocomposite. Results Chem. 2021, 3, 100156.