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Gums and mucilages: versatile excipients for pharmaceutical formulations

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Abstract

Nature has provided us a wide variety of materials to help improve and sustain the health of all living things either directly or indirectly. In recent years there have been important developments in different dosage forms for existing and newly designed drugs and natural products, and semi-synthetic as well as synthetic excipients often need to be used for a variety of purposes. Gums and mucilages are widely used natural materials for conventional and novel dosage forms. These natural materials have advantages over synthetic ones since they are chemically inert, nontoxic, less expensive, biodegradable and widely available. They can also be modified in different ways to obtain tailor-made materials for drug delivery systems and thus can compete with the available synthetic excipients. In this review, we describe the developments in natural gums and mucilages for use in the pharmaceutical sciences.

Keywords: Natural polysaccharide; Natural gum; Pharmaceutical application

1. Introduction

Robbins has stated, "in spite of the problems which have beset the gums market in recent years, the fact remains that in many cases the gums provide a valuable source of income for many poor smallholders or itinerant labourers, either in very poor countries or in the poorest regions rather than more developed countries as such they are important commodities ..." [1]. This remains true today. Tens of thousands of people worldwide, living in regions ranging from semi-arid deserts to rainforests, depend on the collection of gums, resins and latexes in order to provide them with an income. Equally, many millions of people around the world make use of these products in their everyday life [1].

Mother nature has gifted India with great variety of flora and fauna. For centuries man has made effective use of materials of natural origin in the medical and pharmaceutical field. Today, the whole world is increasingly interested in natural drugs and excipients. Natural materials have advantages over synthetic materials because they are non toxic, less expensive and freely available. Furthermore, they can be modified to obtain tailor made materials for drug delivery systems allowing them to compete with the synthetic products that are commercially available. Many kinds of natural gums are used in the food industry and are regarded as safe for human consumption. It should be noted that many 'old' materials are still popular today after almost a century of efforts to replace them. It is usual to strike a balance between economics and performance in the face of commercial realities [2-5].

2. What are gums and mucilages?

Gums are considered to be pathological products formed following injury to the plant or owing to unfavorable conditions, such as drought, by a breakdown of cell walls (extra cellular formation; gummosis) while, mucilages are generally normal products of metabolism, formed within the cell (intracellular formation) and/or are produced without injury to the plant.

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Gums readily dissolve in water, whereas, mucilage form slimy masses.

Gums are pathological products, whereas mucilages are physiological products [6]. Acacia, tragacanth, and guar gum are examples of gums while mucilages are often found in different parts of plants. For example, in the epidermal cells of leaves (senna), in seed coats (linseed, psyllium), roots (marshmallow), barks (slippery elm) and middle lamella (aloe) [7].

Gums and mucilages have certain similarities—both are plant hydrocolloids. They are also translucent amorphous substances and polymers of a monosaccharide or mixed monosaccharides and many of them are combined with uronic acids. Gums and mucilages have similar constituents and on hydrolysis yield a mixture of sugars and uronic acids. Gums and mucilages contain hydrophilic molecules, which can combine with water to form viscous solutions or gels. The nature of the compounds involved influences the properties of different gums. Linear polysaccharides occupy more space and are more viscous than highly branched compounds of the same molecular weight. The branched compounds form gels more easily and are more stable because extensive interaction along the chains is not possible.

3. Disadvantages of synthetic polymers in pharmaceutical sciences

The synthetic polymers have certain disadvantages such as high cost, toxicity, environmental pollution during synthesis, non-renewable sources, side effects, and poor patient compliance.

Acute and chronic adverse effects (skin and eye irritation) have been observed in workers handling the related substances methyl methacrylate and poly-(methyl methacrylate) (PMMA) [8].

Reports of adverse reactions to povidone primarily concern the formation of subcutaneous granulomas at the injection site produced by povidone. There is also evidence that povidone may accumulate in organs following intramuscular injections [9].

Acute oral toxicity studies in animals have indicated that carbomer-934P has a low oral toxicity at a dose of up to 8 g/kg. Carbomer dust is irritating to the eyes, mucous membranes and respiratory tract. So, gloves, eye protection and dust respirator are recommended during handling [10].

Studies in rats have shown that 5% polyvinyl alcohol aqueous solution injected subcutaneously can cause anemia and can infiltrate various organs and tissues [11].

Some disadvantages of biodegradable polymers used in tissue engineering applications are their poor biocompatibility, release of acidic degradation products, poor processing ability and rapid loss of mechanical properties during degradation. It has been shown that poly glycolides, polylactides and their co-polymers have an acceptable biocompatibility but exhibit systemic or local reactions due to acidic degradation products. An initial mild inflammatory response has been reported when using poly-(propylene fumarate) in rat implant studies [12].

4. Advantages of natural gums and mucilages in pharmaceutical sciences

The following are a number of the advantages of natural plant–based materials.

Biodegradable—Naturally available biodegradable polymers are produced by all living organisms. They represent truly renewable source and they have no adverse impact on humans or environmental health (*e.g.*, skin and eye irritation).

Biocompatible and non-toxic—Chemically, nearly all of these plant materials are carbohydrates composed of repeating sugar (monosaccharides) units. Hence, they are non- toxic.

Low cost—it is always cheaper to use natural sources. The production cost is also much lower compared with that for synthetic material. India and many developing countries are dependent on agriculture.

Environmental-friendly processing—Gums and mucilages from different sources are easily collected in different seasons in large quantities due to the simple production processes involved.

Local availability (especially in developing countries)—In developing countries, governments promote the production of plant like guar gum and tragacanth because of the wide applications in a variety of industries.

Better patient tolerance as well as public acceptance—There is less chance of side and adverse effects with natural materials compared with synthetic one. For example, PMMA, povidone.

Edible sources—Most gums and mucilages are obtained from edible sources.

5. Disadvantages of natural gums and mucilages [13-14]

Microbial contamination—The equilibrium moisture content present in the gums and mucilages is normally 10% or more and, structurally, they are carbohydrates and, during production, they are exposed to the external environment and, so there is a chance of microbial contamination. However, this can be prevented by proper handling and the use of preservatives.

Batch to batch variation—Synthetic manufacturing is a controlled procedure with fixed quantities of ingredients, while the production of gums and mucilages is dependent on environmental and seasonal factors.

Uncontrolled rate of hydration—Due to differences in the collection of natural materials at different times, as well as differences in region, species, and climate conditions the percentage of chemical constituents present in a given material may vary. There is a need to develop suitable monographs on available gums and mucilages.

Reduced viscosity on storage—Normally, when gums and mucilages come into contact with water there is an increase in the viscosity of the formulations. Due to the complex nature of gums and mucilages (monosaccharides to polysaccharides and their derivatives), it has been found that after storage there is reduced in viscosity.

6. Classification of gums and mucilages [15-20]

Gums and mucilages are present in high quantities in a varities of plants, animals, seaweeds, fungi and other microbial sources, where they perform a number of structural and metabolic functions; plant sources provide the largest amounts. The different available gums and mucilages can be classified as follows.

6.1. According to the charge

Non-ionic seed gums: guar, locust bean, tamarind, xanthan, amylose, arabinans, cellulose, galactomannans.

Anionic gums: arabic, karaya, tragacant, gellan, agar, algin, carrageenans, pectic acid.

6.2. According to the source

Marine origin/algal (seaweed) gums: agar, carrageenans, alginic acid, laminarin.

Plant origin: (1) shrubs/tree exudates—gum arabica, gum ghatti, gum karaya, gum tragacanth, khaya and albizia gums; (2) seed gums—guar gum, locust bean gum, starch, amylose, cellulose; (3) extracts—pectin, larch gum; (4) tuber and roots—potato starch.

Animal origin: chitin and chitosan, chondroitin sulfate, hyaluronic acid.

Microbial origin (bacterial and fungal): xanthan, dextran, curdian, pullulan, zanflo, emulsan, Baker's yeast glycan, schizophyllan, lentinan, krestin, scleroglucan.

6.3. Semi-synthetic

Starch derivatives—hetastarch, starch acetate, starch phosphates.

Cellulose derivatives—carboxy methyl cellulose (CMC), hydroxy ethylcellulose, hydroxypropyl methylcellulose (HPMC), methylcellulose (MC), microcrystalline cellulose (MCC).

6.4. According to shape

Linear: algins, amylose, cellulose, pectins.

Branched: (1) short branches—xanthan, xylan, galactomanan; (2) branch-on-branch—amylopectin, gum arabic, tragacanth.

6.5. According to manomeric units in chemical structure

Homoglycans—amylose, arabinanas, cellulose; diheteroglycans—algins, carragennans, galactomannans;

tri-heteroglycans—arabinoxylans, gellan, xanthan; tetra-heteroglycans—gum arabic, psyllium seed gum; penta-heteroglycans—ghatti gum, tragacanth.

7. Applications of gums and mucilages

Gums and mucilages of different sources and their derivatives represent a group of polymers widely used in pharmaceutical dosage forms. Various kinds of gums are used in the food industry and are regarded as safe for human consumption.

However, there is growing concern about the safety of pharmaceutical excipients derived from natural sources. Plant gums and exudates are now screened for their use as pharmaceutical adjuvants. Mucilages of different origins are also used in conventional dosage forms of various drugs for their binding, thickening, stabilizing and humidifying properties in medicine. Newer uses of different gums and mucilages in cosmetics and textiles has increased the demand and screening of gums has become an important pharmaceutical area. However, different gums and mucilages used as pharmaceutical adjuvants have stringent specifications, which few natural agents can fulfill. Gums and mucilages have the following applications.

7.1. Applications in the food industry

Gums and mucilages have a variety of applications in the food industry [21]. Different gums have different uses like water retention and stabilizztion (guar and locust bean gum), stabilizers for ice-cream, meat products and instant pudding (carrageenanas), dairy, confectionary and meat products (agar), confectionary, beverages, backed product, and sauces (gum arabic, tragacanth, pectins, alginates and xanthan gum).

7.2. Pharmaceutical applications

Gums and mucilages have a variety of applications in pharmacy. They are used in medicine for their demulcent properties for cough supression. They are ingredients of dental and other adhesives and can be used as bulk laxatives. These hydrophilic polymers are useful as tablet binders, disintegrants, emulsifiers, suspending agents, gelling agents, stabilizing agents, thickening agents, film forming agents in transdermal and periodontal films, buccal tablets as well as sustaining agents in matrix tablets and coating agents in microcapsules including those used for protein delivery. Various gums and mucilages with their common names, biological sources, family and applications are listed in Table 1. Table 2 lists the different applications of gums and mucilages in novel drug delivery systems.

7.3. Industrial uses

Gums used in cosmetics (acacia, tragacanth and karaya gum), textiles (starch, dextrin, cellulose, pectins, and tamarind gum), adhesives (acacia gum, and tragacanth), lithography (gum arabic, tragacanth, and locust bean gum), paints (pectins, hemicellulose, and resins) and paper manufacturer (tamarind, and cellulose).

8. Isolation and purification of gums/mucilages

Plant material is dried in sunlight (preferably) or in an oven at 105˚C to retain its properties unchanged. Generally, chlorophyll or pigments are present in the plant which should be removed before isolating the mucilage. Plant material must be treated with petroleum ether and chloroform (to remove pigments and chlorophyll) and then with distilled water. Care should be taken when drying the final isolated/extracted mucilage. It must be dried at a very low temperature (not more than 50˚C) or in a vacuum. The dried material is stored carefully in desiccators to prevent further moisture uptake or degradation. The general isolation and purification processes for gums and mucilages are shown in Fig. 1.

Baveja *et al*., [22] and Wahi *et al*., [23] reported the following method for the isolation of mucilage. The fresh plant materials were collected washed with water to remove dirt and debris, and dried. Then, the powdered material was soaked in water for 5–6 h, boiled for 30 min, and allows standing 1 h so that all the mucilage was released into the water. The material was then squeezed from an eight muslin bag to remove the

Table 1
Pharmaceutical applications or uses of natural gums and mucilages.

Common name	Botanical name	Family	Pharmaceutical Applications	Reference
Abelmoschus mucilage	*Abelmoschus esculentus*	Malvaceae	Binder in tablets, Sustained release	34, 35
Agar	*Gelidium amansii*	Gelidaceae	Suspending agent, emulsifying agent, gelling agent in suppositories, surgical lubricant, tablet disintegrants, medium for bacterial culture, laxative	36
Albizia gum	*Albizia zygia*	Leguminoseae	Tablet binder	37
Aloe mucilage	*Aloe species*	Liliaceae	Gelling agent, sustained release agent	38
Asario mucilage	*Lepidum sativum*	Cruciferae	Suspending agent, emulsifying agent, controlled release tablet	39, 40
Bavchi mucilage	*Ocimum canum*	Labiatae	Suspending agent, emulsifying agent	39
Carrageenan	*Chondrus cryspus*	Gigarginaceae	Gelling agent, stabilizer in emulsions and suspensions, in toothpaste, demulcent and laxative	41, 42, 43
Cashew gum	*Anacardium occidentale*	Anacardiaceae	Suspending agent	44, 45
Cassia tora	*Cassia tora Linn*	Leguminosae	Binding agent	46
Fenugreek mucilage	*Trigonella foenum graecum*	Leguminoseae	Gelling agent, tablet binder, sustaining agent, emollient and demulcent	22, 47, 48
Guar gum	*Cyamompsis tetraganolobus*	Leguminoseae	Binder, disintegrant, thickening agent, emulsifier, laxative, sustained release agent	49, 50, 51, 52
Gum acacia	*Acacia arabica*	Leguminoseae	Suspending agent, emulsifying agent, binder in tablets, demulcent and emollient in cosmetics	53
Gum ghatti	*Anogeissus latifolia*	Combretaceae	Binder, emulsifier, suspending agent	54
Gum tragacanth	*Astragalus gummifer*	Leguminoseae	Suspending agent, emulsifying agent, demulcent, emollient in cosmetics and sustained release agent	55
Hibiscus mucilage	*Hibiscus esculentus* Linn	Malvaceae	Emulsifying agent, sustained release agent, suspending agent	56, 57
Hibiscus mucilage	*Hibiscus rosasinensis* Linn	Malvaceae	Suspending agent, Sustained release agent	58, 59, 60
Ispagol mucilage	*Plantago psyllium, Plantago ovata*	Plantaginaceae	Cathartic, lubricant, demulcent, laxative, sustaining agent, binder, emulsifying and suspending agent	61, 62, 63, 64, 65
Karaya gum	*Sterculia urens*	Sterculiaceae	Suspending agent, emulsifying agent, dental adhesive, sustaining agent in tablets, bulk laxative	66, 67
Khaya gum	*Khaya grandifolia*	Meliaceae	Binding agent	68
Leucaena seed gum	*Leucaena leucocephata*		Emulsifying agent, suspending agent, binder in tablets, disintegrating agent in tablets	69, 70, 71, 72, 73
Ocimum seed mucilage	*Ocimum gratissimum* L*inn*	Labiatae	Suspending agent, binding agent	74, 75
Pectin	*Citrus aurantium*	Rutaceae	Thickening agent, suspending agent, protective agent	76, 77
Sodium alginate	*Macrocytis pyrifera*	Lessoniaceae	Suspending agent, gelation for dental films, stabilizer, sustained release agent, tablet coating	79, 80, 81, 82, 83
Satavari mucilage	*Asparagus racemosus*	Aapocynaceae	Binding agent and sustaining agent in tablets	78

Table 1
Pharmaceutical applications or uses of natural gums and mucilages (Continued).

Common name	Botanical name	Family	Pharmaceutical applications	Reference
Tamarind seed polysaccharide	*Tamarindus indica*	Leguminoseae	Binding agent, emulsifier, suspending agent, sustaining agent	84
Xanthan gum	*Xanthomonas lempestris*	---	Suspending agent, emulsifier, stabilizer in toothpaste and ointments, sustained release agent	66, 85
Gellan gum	*Pseudomonas elodea*	---	Disintegrating agent,	86

Table 2
Applications of gums and mucilages in NDDS.

Common name	Botanical name	Family	Pharmaceutical applications	Reference
Acacia	*Acacia Senegal*	Leguminosae	Osmotic drug delivery	87, 88
Bhara gum	*Terminalia bellerica* roxb	Combretaceae	Microencapsulation	89
Chitosan	—	—	Colonspecific drug delivery, microspheres, carrier for protein as nanoparticles	90, 91
Cordia gum	*Cordia obliqua* willed	Boraginaecae	Novel oral sustainedrelease matrix forming agent in tablets	92
Cactus mucilage	*Opuntia ficus-indica*	—	Gelling agent in sustained drug delivery	93
Guar gum	*Cyamompsis tetraganolobus*	Leguminoseae	Colontargeted drug delivery, cross-linked microspheres	94, 95, 96
Gellan gum	*Pseudomonas elodea*	—	Ophthalmic drug delivery, sustaining agent, beads, hydrogels, floating in-situ gelling, controlledrelease beads	97, 98, 99, 100, 101, 102
Hakea	Hakea gibbosa	—	Sustainedrelease and peptide mucoadhesive for buccal delivery	103, 104
Ispagol	*Plantago psyllium, Plantago ovata*	Plantaginaceae	Hydrogels, colon drug delivery, gastroretentive drug delivery	105, 106, 107, 108
Karaya gum	*Sterculia urens*	Sterculiaceae	Mucoadhesive and buccoadhesive	109
Locust bean gum	*Ceratania siliqua*	Leguminoseae	Controlledrelease agent	110
Mucuna gum	*Mucuna flagillepes*	Papillionaceae	Microspheres	111
Okra	*Hibiscus esculentus*	Malvaceae	Hydrophilic matrix for controlled release drug delivery	112
Pectin	*Citrus aurantium*	Rutaceae	Beads, floating beads, colon drug delivery, pelletization by extrusion/spheronization, microparticulate delivery, transdermal delivery, Iontophoresis, hydrogels	113, 114, 115, 116, 117, 118, 119, 120
Sodium alginate	*Macrocytis pyrifera*	Lessoniaceae	Bioadhvesive microspheres, nanoparticles, microencapsulation	121
Tamarind	*Tamarindus indica*	Leguminoseae	Hydrogels, mucoadhesive drug delivery for ocular purposes, spheroids, nasal drug delivery	122, 123
Xanthan gum	*Xanthomonas lempestris*	—	Pellets, controlled drug delivery system	124, 125

marc from the solution. Following this, three volumes of acetone was added to the filtrate to precipitate the mucilage. The mucilage was separated, dried in an oven at a temperature less than 50˚C, and the dried powder was passed through a No. 80 sieve and stored in a desiccator until required. The isolated mucilage from the plant was subjected to some preliminary confirmative testing. Table 3 shows the preliminary confirmative test for dried mucilage [6, 15, 16].

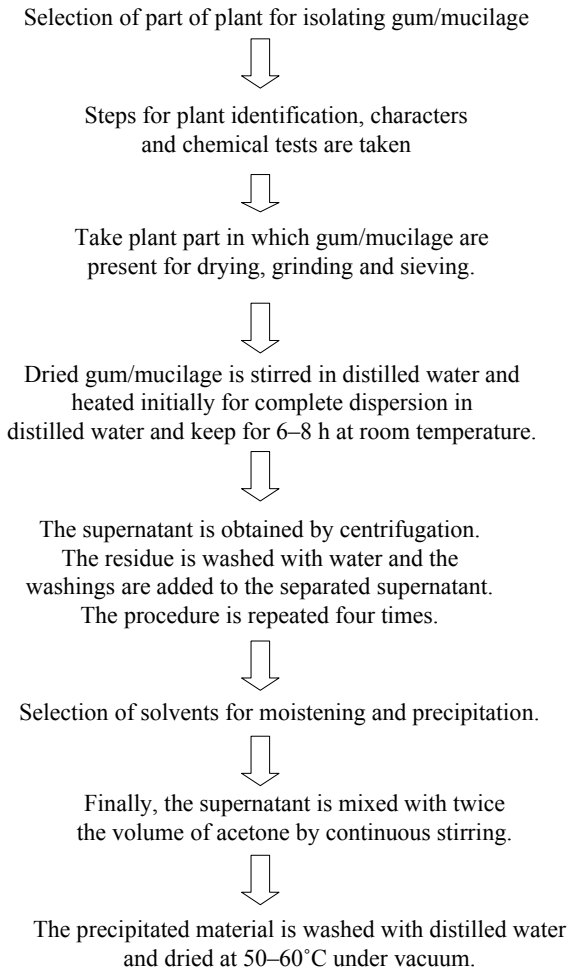

Fig. 1. General isolation/extraction procedure for mucilages.

Extraction is one of the most crucial procedures to achieve complete recovery of target compounds from plants. Recently, microwave energy has started to be used for the extraction of phytoconstituents from plants [24]. It is a simple, fast, clean, eco-friendly and efficient method and saves energy, fuel and electricity. Microwave extraction follows the same principle as maceration or percolation, but the speed of breaking up of the plant cells and tissues is much higher. Microwave assisted extraction methods require a shorter time and less solvent, and provide a higher extraction rate and better products at a lower cost. Plant material is powdered in a mechanical blender for 5 m and then soaked in distilled water for 24 h in a 1000 ml beaker. It is kept in a microwave oven along with a glass tube to prevent bumping when subjected to microwave irradiation. The beaker is removed from the oven and allowed to stand for 2 h to allow the mucilage to be released into the water. It is then processed in a similar way to the conventional procedure, weighed and stored.

9. Characterization / Standardization of gums and mucilages

A suitable strategy is required to save money and time. Over-characterization is not desirable, because excessive use of time and resources could actually delay

Table 3
Preliminary confirmative tests for dried mucilage powder.

Test	Observation	Inferences
Molisch's test: (100 mg dried mucilage powder + Molisch's reagent + conc. H_2SO_4 on the side of a test tube)	Violet green color observed at the junction of the two layers	Carbohydrate present
Ruthenium test: Take a small quantity of dried mucilage powder, mount it on a slide with ruthenium red solution, and observe it under microscope.	Pink color develops	Mucilage present
Iodine test: 10 0mg dried mucilage powder + 1 ml 0.2 N iodine solution.	No color observed in solution	Polysaccharides present (starch is absent)
Enzyme test: dissolve 100 mg dried mucilage powder in 20 ml-distilled water; add 0.5 ml of benzidine in alcohol (90%). Shake and allow to stand for few minutes	No blue color produced	Enzyme absent (Distinction between dried mucilage and acacia)

the launch of innovative excipients. The characterization of gums and mucilages is initially achieved by only a multiple-technique approach [25]. For excipient analysis, analytical techniques can be classified according to the type of information generated.

Structural—Gums and mucilages are polysaccharides and contain sugars. So, confirmation of the different sugars is carried out by chromatography and structure elucidation can be carried out by NMR and mass spectroscopy.

Purity—To determine the purity of the selected gum and mucilage, tests for alkaloids, glycosides, carbohydrates, flavanoids, steroids, amino acids, terpenes, saponins, oils and fats, and tannins and phenols are carried out.

Impurity profile—Testing for impurities must be carried out using suitable analytical techniques.

Physico-chemical properties—Color, odor, shape, taste, touch, texture, solubility, pH, swelling index, loss on drying, hygroscopic nature, angle of repose, bulk and true densities, porosity and surface tension. Different ash values are also estimated. The microbial load and presence of specific pathogens are also determined. *In vitro* cytotoxicity is also determined. Gums and mucilages are highly viscous in nature. So, the rheological properties of excipients are important criteria for deciding their commercial use. The flow behavior of the samples is determined.

Toxicity—The acute toxicity of gums and mucilages are determined by the followings fixed-dose method as per OECD guideline No. 425. A sub-acute toxicity study, determination of the LD50 etc., is carried out in rats and guinepigs of both sexes.

Once analysis is complete, determination of the structure, composition and impurity profile enables a scientific dossier to be prepared describing the excipient. This information is of value for the regulatory dossier of the final pharmaceutical product that would contain the given excipient.

Finally, gums and mucialges are added to pharmaceutical formulations. So a compatibility study is important. The compatibility studies of gum/mucilage/drugs are performed using spectrophotometry/FTIR/DSC.

10. Pharmacopoeial standard specifications of gums and mucilages

Different pharmacopoeias, like USP, PhEur, and JP give pharmacopoeial standards for specific gums [26]. These are shown in Table 4.

11. Reasons for developing new excipients

For a number of reasons there has been an increase in interest in the development of new excipients/diluents.

Some drugs show incompatibilities with many of the current range of excipients. For example, atenolol-PVP, atenolol-mg-stearate [27]. One of the more common drug-excipient incompatibilities is the reaction between aldehydic sugars, such as lactose and primary and secondary amines, leading to the formation of Schiff bases. These complex series of reactions lead to browning and discoloration of the dosage form. Despite being a carrier of choice for dry powder aerosol formulations, lactose may need to be replaced with a different carrier, such as mannitol or sucrose, when formulating primary and secondary amines.

Mg-stearate is incompatible with aspirin, some vitamins and most alkaloidal salts [28].

There is a need for excipients that will allow faster manufacturing of formulations. For example, at the present time, in tablet dosage forms, new excipients having better compressibility at very high compression speeds are needed. Today, it is not unheard of to have tableting equipment compressing 8000 to 10000 tablets per min. It is critical under these conditions to have an exceptionally efficient flowing granulation/powder blend. Many sugar-based excipients, such as maltose, mannitol, and sorbitol are not compressible in their natural state and need to be modified for use in direct compression tableting.

Some future developments may require new delivery systems. For example, new drug delivery systems for oral administration of biotechnology products need new excipients which will avoid the inconvenience of multiple daily injections. Progress in the development of peptides as therapeutic drugs has been impeded in part by their rapid excretion, resulting in short

circulating lifetimes. This has generated considerable interest in improving the duration of action of drugs through conjugation with the water-soluble, biocompatible excipient, poly (ethylene glycol). Such conjugates have reduced enzymatic degradation rates and lengthened circulating lifetimes compared with the native compounds. There are six FDA-approved PEGylated products on the market, vouching for the safety and commercial viability of this technology. Other novel lipophilic carbohydrate excipients, termed oligosaccharide ester derivatives (OEDs), have been used to modify the pharmacokinetic profiles of drugs. This technology is quite flexible, offering the ability to formulate drug molecules with modified-release characteristics and improved bioavailability. In other areas of technology, selected carbohydrate excipient, such as trehalose and sucrose to stabilize molecules in the dry state, thereby preventing their physical and chemical degradation at ambient temperatures and above. These patent-protected drug delivery technologies are suited to the delivery of macromolecules, such as proteins and peptides by the pulmonary, oral, and injectable routes.

Drug targeting systems, like liposome delivery systems, need newer excipient, because the existing excipients for liposomes are too expensive.

12. Modification of existing gums and mucilages

It should be noted that many ‘‘old’’ materials compete successfully today after almost a century of efforts to replace them. It is the usual balance of economics and performance that determines the commercial realities. Natural gums have been modified to overcome certain drawbacks, like uncontrolled rate of hydration, thickening, drop in viscosity on storage, and microbial contamination [29].

Since the implementation of polymeric materials in the field of pharmaceutical technology, numerous attempts have been made to modify their physical and chemical properties, and thus, their potential applicability in various areas of drug formulation.

Various methods are available to modify the state of molecular interaction between polymers. Basically, two methods are available as the physical method and chemical method.

Physical method—a molecular interaction between polymers can be achieved by exposure to dry heat, saturated steam, microwave technology, UV [30-31], and gamma radiation [32].

Chemical method—polymers are treated with

Table 4
Pharmacopoeial specifications for gums.

Excipient	Test	Pharmacopeia
Acacia	Microbial limit, ash values,	USP, JP, PhEur
Alginic acid	Microbial limit, pH, loss on drying	USP, PhEur
Carrageenan	Solubility, viscosity, loss on drying, ash value	USP
Dextrin	Loss on drying, residue on ignition, reducing sugars	USP, BP, JP
Gelatin	Isoelectric point, microbial limit, residue on ignition, loss on drying, total ash, jelly strength	USP, JP, PhEur
Guar gum	pH, microbial contamination, apparent viscosity, loss on drying, ash, galactomannans, organic volatile impurities,	USP, PhEur
Lecithin	Water, arsenic, lead, acid value, heavy metals	USP
Sodium alginate	Microbial limit, appearance of solution, loss on drying, ash, heavy metals	USP, PhEur
Tragacanth	Microbial limits, flow time, lead, acacia and other soluble gums, heavy metals	USP, JP, PhEur
Xanthan gum	pH, viscosity, microbial limits, loss on drying, ash, heavy metals, organic volatile impurities	USP, PhEur
Gellan gum	pH, microbial limit, loss on drying, moisture content, specific gravity, solubility, bulk density	USP

chemicals like aldehydes, epichlorhydrin, borax or glutaraldehyde. Temperature is one of the most favorable methods of cross-linking because it avoids both the application of harsh chemical materials for large-scale production and the diversity of equipment and methods used in their application [33]. Table 5 shows examples of modified gums and mucilages.

13. Precautions

Since they (gums and mucilages) deteriorate when stored, especially in warm weather preservatives are added, such as solutions of formaldehyde (10 min per pint) or benzoic acid (10 grains per pint).

14. Conclusion

Natural gums are promising biodegradable polymeric materials. Many studies has been carried out in fields including food technology and pharmaceuticals using gums and mucilages. It is clear that gums and mucilages have many advantages over synthetic materials. Various applications of gums and mucilages have been established in the field of pharmaceuticals. However, there is a need to develop other natural sources as well as with modifying existing natural materials for the formulation of novel drug delivery systems, biotechnological applications and other delivery systems. Therefore, in the years to come, there will be continued interest in natural gums and their modifications aimed at the development of better materials for drug delivery systems.

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Table 5
Examples of modified gums with their applications.

Gums and mucilage	Modification technique	Application	References
Karaya gum	Heat Treatment at various temperatures in a hot air oven.	Disintegrating agent	126, 127
Agar and Guar gum	Heat Treatment at various temperatures in a hot air oven along with co-grinding of both materials.	Disintegrating agent	128
Hypochlorite potato starch	Chemical modification of potato starch carried out in presence of hypochloride.	Disintegrating agent	129
Tragacanth	Chemical modification of tragacanth using epichlorhydrine.	Disintegrating agent	130
Acacia gum	Chemical modification of acacia gum using epichlorhydrine.	Disintegrating agent	131
Guar gum	Chemical modification of guar gum	Disintegrating agent	132
Cross-linked amylose	Chemical modification of amylase by substituting it in a one step reaction.	Disintegrating and binding agent	133
Cross-linked cellulse	Chemical modification of cellulose by epichlorhydrine.	Disintegrating and binding agent	134
Polyalkylamine	Chemical modification of polyalkylamine.	Disintegrating agent	135
Cyclodextrin	Physical modification - co-drying of micro crystalline cellulose with cyclodextrin	Disintegrating agent	136
Starch	Physico-chemical treatment of to starch for modification	Disintegrating and binding agent	137
Sesbania gum	Chemical modification of *Sesbania* gum with tartaric acid for a sustained release formulation and chemical modification of gum with acetone: chloroform mixture for gelling agent	Sustained release formulation, gelling agent	138, 139
Guar gum	Chemical modification of guar gum with glutaraldehyde for colonic delivery, chemical modification using isopropanol as a filmcoating material	Colonic delivery, film coating, hydrogel	140,141, 142, 143, 144
Tamarind powder	Chemical modification of tamarind powder using epichlorohydrin for a sustainedrelease formulation and partial degradation of β-galactosidase for rectal drug delivery.	Sustainedrelease formulation, rectal drug delivery	145, 146
Psyllium	Chemical modification of psyllium was carried out to form *N*-hydroxymethylacrylamide based hydrogels, chemical modification with tartaric acid.	N-hydroxymethylacrylamide based hydrogels, oral insulin drug delivery	147, 148, 149, 150
Okra fruits (pods) of Hibiscus esculentus	Chemical modification with acrylamide synthesis	Controlled drug delivery	151
Ipomoea dasysperma, Ipomoea hederacea, and Ipomoea palmata	Chemical modification of ipomoea with poly(acrylonitrile) grafted drug delivery	Poly(acrylonitrile) grafted drug delivery	152
Pectins	Chemical modification of pectin with acetyl chloride in ethanol for modified drug delivery, chemical modification with ethanolamine for hydrogels and chemical modification of pectin for colonic drug delivery.	Modified drug delivery. hydrogels, colonic drug delivery	153, 154, 155

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