

<u>Pharmaceutical medications and natural therapeutic products for</u> <u>managing diabetes mellitus</u>

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Introduction:

Diabetes mellitus is a prevalent and long-lasting disease that affects children and adolescents in both developed and developing countries worldwide. Diabetes mellitus encompasses two primary types: type I and type II. The former refers to an autoimmune condition that results in malfunction of the pancreatic β -cells, resulting to insufficient generation of insulin. The latter condition is caused by decreased responsiveness to insulin in the specific tissues (such as skeletal muscle, liver, and adipose tissue), followed by inadequate production of insulin. Both scenarios lead to a consistent increase in blood sugar levels, known as sustained hyperglycemia. Prolonged uncontrolled hyperglycemia results in the progressive deterioration of the blood vessels that provide nourishment to the organs of the body, resulting in damage to the heart, eyes, kidneys, and nervous system. Macrovascular (atherosclerotic) and microvascular (retinopathy and nephropathy) problems are the primary reasons for illness and death in individuals with diabetes. Diabetes is a complex condition that encompasses a group of metabolic problems, rather than being a single disease (Zaid, et al.2016).

Pharmaceutical drugs and natural therapeutic materials have gained significant attention for their potential in managing diabetes among the wide range of therapeutic approaches available. Pharmaceutical drugs, such as insulin analogues, oral hypoglycemic medicines, and injectable therapy, have been essential in the treatment of diabetes mellitus for a long time. These drugs achieve their effects through different mechanisms, including increasing insulin secretion, boosting insulin sensitivity, decreasing hepatic glucose synthesis, and delaying carbohydrate absorption. Although pharmaceutical drugs have shown efficiency in controlling blood sugar levels and reducing the chances of problems, there are ongoing worries about their safety, acceptability, and cost-effectiveness. As a result, researchers are investigating alternate therapy methods (Zhao, et al.2020).

Although synthetic medications have made significant advancements, herbal medicine is widely used in both developed and developing nations. Moreover, the global demand for herbal medicine preparations has significantly risen in recent decades, likely due to the enduring effectiveness of this form of therapy. Natural goods contain a wide range of bioactive components, such as polyphenols, flavonoids, alkaloids, and terpenoids. These chemicals have many ways of affecting the body, such as increasing insulin secretion, improving glucose uptake, regulating inflammatory pathways, and safeguarding pancreatic β cells. Furthermore, herbal medicines are comparatively affordable and are perceived to be less risky than synthetic treatments. However, it is important to exercise caution prior to administering any herbal medication treatment (Verma, et al.2018).

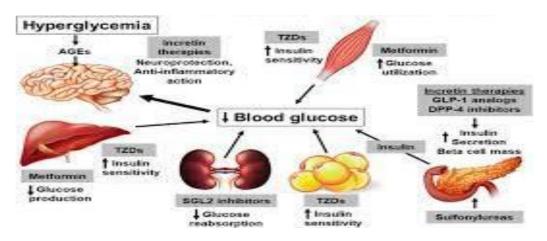


4 Mechanisms of Action in Diabetes Management:

Pharmaceutical Medications:

Given the complex nature of the development of diabetes mellitus, effectively managing and treating the condition involves considering the individual circumstances of patients. This includes implementing self-management strategies such as modifying diet, engaging in appropriate exercise, closely monitoring glucose levels, assessing mood, and using medication as necessary. The medications used for treating diabetes mostly include insulin, insulin analogues, and non-insulin hypoglycemic drugs. Non-insulin hypoglycemic drugs can be further categorized as insulin sensitizers, insulin secretagogues, glucose regulators, and gene therapy (Zhao, et al.2020).

- **Insulin Analogues**: imitate natural insulin by attaching to insulin receptors on specific cells, which helps in the absorption of glucose into muscle and fat tissues, and reduces the creation of glucose by the liver. Rapid-acting analogues facilitate the regulation of glucose levels after meals, whereas long-acting analogues offer continuous insulin coverage (Petersen, & Shulman, 2018).
- Oral hypoglycemic Agents: medicines, including sulfonylureas, biguanides, thiazolidinediones, and dipeptidyl peptidase-4 (DPP-4) inhibitors, act on distinct areas of glucose metabolism. For instance, sulfonylureas enhance the release of insulin from pancreatic β-cells, whilst biguanides enhance the ability of peripheral tissues to respond to insulin and inhibit the production of glucose by the liver.
- Injectable Therapies: such as glucagon-like peptide-1 (GLP-1) receptor agonists and sodium-glucose cotransporter-2 (SGLT-2) inhibitors, affect the incretin and renal glucose reabsorption pathways, respectively. GLP-1 receptor agonists increase insulin production in response to glucose levels, slow down the emptying of the stomach, and decrease appetite. On the other hand, SGLT-2 inhibitors increase the excretion of glucose through urine and decrease the reabsorption of glucose by the kidneys (Anderson, 2020).





- Safety:

Although pharmacological drugs are typically efficacious, they may be linked to specific safety issues such as hypoglycemia, weight gain, gastrointestinal side effects, and cardiovascular risks. It is crucial to closely monitor and make personalized adjustments to treatment in order to reduce these risks and get the best possible safety outcomes.

- Natural Therapeutic Products:
 - **Botanicals and Dietary Supplements:** Natural products derived from botanical sources, such as bitter melon, fenugreek, cinnamon, and Gymnema sylvestre, contain bioactive compounds with hypoglycemic properties. These compounds may act through multiple mechanisms, including enhancing insulin secretion, improving insulin sensitivity, and inhibiting carbohydrate digestion and absorption. Medicinal plants or plant-based medicine have been utilized globally in a costefficient manner for the prevention and/or treatment of diabetes. Indeed, numerous underdeveloped nations depend on herbal medicine to provide treatment for individuals suffering from diabetes and other ailments. Many medications currently in use are structurally derived from natural chemicals found in traditional medicinal plants. Metformin, an anti-hyperglycemic medication used for diabetic treatment, can be attributed to the historical use of Galega officinalis, a traditional remedy for diabetes. Some of the frequently utilized medicinal plants and vitamins with hypoglycemic properties that can enhance the immune system and regulate blood sugar levels in humans are Allium sativum (garlic), Momordica charantia (Bitter Melon), Hibiscus sabdariffa L. (Roselle Plant), Zingiber officinale Rosc (Ginger), and Vitamins C, D, and E. Due to their easy accessibility, affordability, and efficacy in managing diabetes, both developing and affluent nations rely on medicinal plants to fulfill their healthcare requirements (Yedjou, et al.2023).

1. Allium sativum:

Allium sativum is the taxonomic designation for garlic, a member of the Amaryllidaceae family. Garlic possesses numerous health advantages, including the ability to reduce cholesterol levels, enhance blood pressure, and strengthen the immune system. Studies have demonstrated that garlic may offer advantages for those with diabetes. Garlic contains several important phytochemicals, including allicin, which is responsible for its strong smell (Melino, et al.2019). Other sulfur compounds found in garlic, such as diallyl disulfide and diallyl trisulfide, have beneficial properties like antioxidant, anti-inflammatory, anti-diabetes, and anti-cancer effects. Garlic also contains flavonoids like quercetin and kaempferol, which are known for their antioxidant properties. Additionally, garlic contains saponins, which are natural detergents that can help lower cholesterol levels. Lastly, garlic contains fructans, a type of carbohydrate that can act as prebiotic candidates. Garlic has numerous health benefits, such as its ability to reduce inflammation,



modulate the immune system, protect the heart, lower cholesterol levels, decrease blood sugar, function as an antioxidant, fight against bacteria, fungi, and viruses, and prevent the growth of cancer cells (Shang, et al.2019).

2. Momordica charantia:

Momordica charantia, commonly referred to as bitter melon, is utilized as a supplementary or alternative therapeutic approach for managing DM in both developing and affluent nations. It contains chemicals that have a significant impact on controlling and reducing blood glucose levels in persons with diabetes mellitus. It controls and decreases blood glucose levels in individuals with diabetes by enhancing insulin sensitivity and lowering glucose synthesis in the liver. Bitter melon has several health benefits due to its anti-inflammatory, immunomodulatory, hypolipidemic, hypoglycemic, antioxidant, antifungal, antibacterial, anticancer, and antiviral properties (Alam, et al.2015).

3. Hibiscus sabdariffa L. (Roselle):

Its long history in traditional medicine is attributed to its abundant presence of pharmacologically active chemicals and notable therapeutic capabilities. Hibiscus sabdariffa L. (roselle) has been widely claimed to have numerous health benefits, including its ability to lower blood pressure, reduce inflammation, decrease body fat mass, modulate the immune system, lower blood sugar levels, act as an antioxidant, lower cholesterol levels, prevent cancer, and alleviate dry mouth (Patel, 2014). The medicinal benefits of Hibiscus sabdariffa L. (roselle) are attributed to the presence of bioactive and functional components, including phenolic acids, flavonoids, anthocyanins, organic acids, and dietary fiber (Riaz, & Chopra, 2018).

- Herbal Remedies: Traditional herbal remedies, such as ginseng, berberine, and Momordica charantia (bitter melon), have been used for centuries in various cultures for diabetes management. These herbs may exert their effects by modulating insulin signaling pathways, activating AMP-activated protein kinase (AMPK), and enhancing glucose uptake into cells (Kasole, et al.2019).
- **Nutritional Supplements:** Nutritional supplements, including omega-3 fatty acids, chromium, and alpha-lipoic acid, have been studied for their potential in improving glycemic control and reducing insulin resistance. These supplements may influence cellular metabolism, oxidative stress, and inflammation, thereby impacting glucose homeostasis (Kasole, et al.2019).
- Safety:

Natural therapeutic products are often perceived as safer alternatives to pharmaceutical medications due to their natural origins. However, they may still pose safety risks, including potential herb-drug interactions, variability in potency and purity, and lack of standardized dosing regimens. Adverse effects such as gastrointestinal upset, allergic reactions, and hepatotoxicity have been reported with certain



natural products, underscoring the importance of caution and supervision when using these remedies.

- **4** Drug Interactions:
- Pharmacokinetic interactions:
 - Metabolism and Elimination: Numerous pharmaceutical drugs utilized in the management of diabetes undergo metabolic processes facilitated by cytochrome P450 (CYP) enzymes located in the liver. Therapeutic goods derived from nature that contain bioactive chemicals, such as flavonoids, alkaloids, and terpenoids, have the ability to either inhibit or stimulate these enzymes, resulting in changes to the way drugs are processed and removed from the body. St. John's wort, a frequently used herbal treatment, is recognized for its ability to stimulate CYP3A4, which can potentially decrease the effectiveness of specific oral hypoglycemic medications and insulin. Natural medicinal materials can potentially impact the absorption and distribution of drugs by modifying gastrointestinal motility, intestinal permeability, and drug transporter activity. For example, using dietary fiber supplements like psyllium and guar gum might slow down the absorption of oral drugs, which can result in decreased effectiveness or inconsistent blood glucose levels (Blahova, et al.2021).
- Pharmacodynamic interactions:
 - **Hypoglycemia:** is a major worry when it comes to drug interactions in diabetes management. Pharmaceutical drugs like sulfonylureas and insulin can reduce blood glucose levels. When used together with natural medicinal remedies that also lower blood sugar, like bitter melon and fenugreek, this can raise the risk of hypoglycemia.
 - **Cardiovascular Effects:** Certain natural medicinal agents, like ginseng and ginkgo biloba, can potentially impact the cardiovascular system by regulating blood pressure and inhibiting platelet aggregation. Simultaneous use of antihypertensive medicines or anticoagulants used in diabetes treatment may result in additional effects or an increased risk of bleeding (Sheneni, & Shaibu, 2023).
 - **4** Side Effects and Adverse Reactions:
- Pharmaceutical Medications:

Pharmaceutical drugs utilized for diabetes control can be linked to a variety of adverse effects, which differ in occurrence and intensity based on the medication and specific patient characteristics. Typical adverse effects of insulin therapy encompass hypoglycemia, weight gain, injection site responses, and allergic reactions. Oral hypoglycemic medications, such as sulfonylureas and thiazolidinediones, can lead to hypoglycemia, gastrointestinal discomfort, fluid retention, and increased body weight. Injectable treatments, such as GLP-1 receptor agonists and SGLT-2 inhibitors, might cause gastrointestinal problems, reactions at the injection site, and urinary tract infections. Although the majority of patients can



handle pharmaceutical prescriptions without any issues, certain individuals may encounter negative effects that have a significant impact on their overall well-being and ability to follow their treatment plan. Hypoglycemia, specifically, can be a notable issue, as it can result in symptoms such as perspiration, tremors, cognitive disorientation, and in severe instances, unconsciousness. Weight gain caused by specific drugs might provide challenges for individuals with diabetes who are working towards their weight management objectives (Blahova, et al.2021).

• Natural Therapeutic Products:

While natural medicinal solutions used in diabetes control are commonly seen as safer alternatives to pharmaceutical drugs, they can nevertheless have side effects and severe responses. Typical adverse effects of natural goods may encompass gastrointestinal disturbance, allergic responses, and interactions with concurrent drugs. For instance, the consumption of bitter melon may result in gastrointestinal discomfort and hypoglycemia, whilst high quantities of cinnamon supplements can potentially cause allergic reactions and liver toxicity. Many patients, especially those looking for supplementary or alternative treatments for diabetes, often handle natural therapeutic materials well. Nevertheless, it is important to note that people's reactions to these products might differ, and certain persons may encounter negative consequences that require them to stop using the product or modify their dosage. Healthcare practitioners must educate patients about the potential hazards and advantages of natural products and closely monitor for any negative responses during treatment.

4 Personalized Medicine Approaches:

• Genetic Testing:

Genetic testing can identify genetic variants associated with diabetes risk, treatment response, and susceptibility to complications. Understanding a patient's genetic predisposition can inform treatment decisions and personalize interventions. Pharmacogenomic testing can identify genetic variations that affect drug metabolism, response, and toxicity. This information can guide the selection of appropriate pharmaceutical medications and dosages based on an individual's genetic profile. Additionally, genetic testing can identify patients who may benefit from specific natural therapeutic products targeting genetic pathways implicated in diabetes pathophysiology (Sugandh, et al.2023).

• Biomarker Analysis:

Biomarker analysis involves assessing biological markers, such as HbA1c levels, lipid profiles, and inflammatory markers, to stratify patients based on disease severity, progression, and treatment response. Biomarkers can provide insights into underlying disease mechanisms and predict treatment outcomes. Biomarker analysis can guide the selection of pharmaceutical medications and natural therapeutic products tailored to an individual's disease phenotype and metabolic profile. For example, patients with



insulin resistance may benefit from medications targeting insulin sensitivity, while those with elevated inflammatory markers may benefit from anti-inflammatory natural products.

• Individualized Treatment Plans:

Individualized treatment plans take into account a patient's unique characteristics, preferences, and goals to develop personalized interventions. These plans integrate information from genetic testing, biomarker analysis, clinical assessments, and patient-reported outcomes to tailor treatment regimens. Individualized treatment plans can optimize treatment outcomes by selecting the most effective and well-tolerated pharmaceutical medications and natural therapeutic products for each patient. For example, patients with specific genetic variants associated with metformin response may benefit from this medication as a first-line therapy, while others may require alternative agents. Similarly, patients with biomarker profiles indicative of insulin resistance or beta-cell dysfunction may benefit from targeted interventions, such as GLP-1 receptor agonists or botanicals with insulin-sensitizing properties.

4 Patient Preferences and Compliance:

• Cultural beliefs and traditions:

Patients' perceptions of sickness and therapy might be greatly influenced by their cultural beliefs and traditions. Certain cultural groups may exhibit a predilection for natural medicines derived from old medical traditions, whereas others may place their faith in contemporary pharmaceutical medications. Healthcare practitioners should actively participate in culturally sensitive communication and demonstrate respect for patients' cultural values when discussing treatment alternatives. Gaining insight into cultural norms and preferences can enhance trust and promote cooperation within the therapeutic alliance (Sari, et al.2022).

• Perceived effectiveness and safety:

Patients' opinions of the effectiveness and safety of pharmaceutical pharmaceuticals and natural medicinal products can differ depending on their personal experiences, family background, and cultural factors. Presenting empirical data on the efficacy and security of treatment alternatives can assist patients in making well-informed choices. By emphasizing the advantages and possible drawbacks of each treatment method, we can correct misunderstandings and increase the likelihood of patients accepting the treatment.

• Expense and availability:

The factors that strongly influence treatment adherence are the cost and accessibility. Patients may encounter financial limitations or experience limited availability of healthcare services, which can impact their capacity to procure pharmaceuticals or natural products. Healthcare professionals should engage in transparent discussions regarding treatment costs and actively seek various resources, including insurance coverage, prescription assistance programs, and community health services, in order to address and



minimize financial obstacles. Suggesting cost-effective and easily obtainable treatment alternatives might enhance compliance and health results (Alowayesh, et al.2022).

• Convenience and user-friendliness:

Patient adherence is significantly influenced by the convenience and simplicity of the treatment. Patients may have a preference for medications or items that are easy to use, have few adverse effects, and can be incorporated into their regular schedules. Enhancing treatment adherence can be achieved by providing flexible dose schedules, streamlined treatment regimens, and user-friendly administration techniques. Offering explicit guidelines and supplementary resources can enable individuals with diabetes to effectively control their condition.

• Enhancing patient knowledge and autonomy:

Providing patients with education is crucial for promoting their compliance with treatment and development of skills to control their health. Providing patients with information on the significance of adhering to medication, regularly testing blood glucose levels, and adopting good lifestyle behaviors can empower them to actively participate in managing their diabetes. Fostering transparent dialogue, acknowledging apprehensions, and offering continuous assistance can cultivate patient assurance and fortitude in handling their ailment (Sallay, et al.2021).

Conclusion:

The use of personalized medicine strategies has great potential for transforming diabetes management by customizing interventions based on the unique characteristics of each patient. Healthcare professionals can enhance treatment outcomes and enhance patient care by utilizing genetic testing, biomarker analysis, and personalized treatment programs. This applies to both pharmaceutical drugs and natural therapeutic solutions. These methods enable the detection of genetic variations, metabolic indicators, and illness characteristics that affect the response to treatment and the likelihood of experiencing problems. Healthcare practitioners can empower people with diabetes by adopting personalized medicine, which involves providing tailored interventions that specifically address their individual requirements, preferences, and goals.



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