

A STUDY OF SYNTHESIS AND BIO-ASSESSMENT OF ANTICANCER-BASED COMPOUNDS

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ABSTRACT

An in-depth analysis of the bio-assessment and synthesis of compounds with dual functionality—potential anticancer activity and Diacylglycerol O-acyltransferase 1 (DGAT1) inhibition—is presented in this paper. The emergence of DGAT1 as a potential therapeutic target in metabolic disorders has prompted the development and manufacturing of drugs with inhibitory effects. DGAT1 is an important enzyme in lipid metabolism. Research into compounds with dual actions beyond DGAT1 inhibition is motivated by the study's recognition of the urgent need for novel anticancer medicines. In order to create molecules that inhibit DGAT1 and have anticancer properties, the synthesis step entails a thorough investigation of several synthetic techniques. Molecular structures are designed and modified according to principles of medicinal chemistry with the goal of achieving the best inhibitory effects against DGAT1 while also including aspects that could be anticancer. In order to guarantee scalability and reproducibility, the research follows complex synthetic pathways, optimizing chemical reactions and circumstances. The next step is bio-assessment, when the produced chemicals are tested extensively to determine their safety and effectiveness. The chemicals' cytotoxicity, antiproliferative effects, and influence on cellular processes can be better understood by in vitro experiments that use cell lines that are indicative of metabolic diseases and different forms of cancer. Understanding the systemic effects, metabolism, and possible toxicities is the goal of the bio-assessment, which includes in vivo investigations conducted on preclinical animal models. In light of the chemicals' possible use as immunomodulatory agents in cancer treatment plans, we investigate how they interact with the immune system.

KEYWORDS: Bio-Assessment, Anticancer-Based Compounds, anticancer activity, DGAT1 inhibition, antiproliferative effects.

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INTRODUCTION

We get protein, fat, and carbs from the food we consume. Enzymes are chemicals that the body secretes once food reaches the digestive tract. Enzymes like these break down macromolecules of sugar, lipid, and protein into their component amino acids, fatty acids, and carbs, respectively. Proteins, carbohydrates, and lipids are all potential sources of energy for our bodies. The absorption of these substances into the bloodstream carries them to the cells. Various biochemical enzymes control the metabolic processes that take place when these substances enter the cells. A live thing's metabolic process entails cellular chemical reactions. The metabolic process keeps cells and creatures alive by transforming food into energy. Because they conduct chemical reactions in the correct direction, enzymes are crucial to metabolism. Metabolic pathways are the series of events that cause certain chemical reactions to take place. Metabolic pathways are well-documented in biochemistry. Amino acid biosynthesis, purine biosynthesis, fatty acid metabolism, inositol metabolism, glucuronate metabolism, histidine metabolism, aromatic amino acid biosynthesis, citric acid cycle, urea cycle, and many more key metabolic pathways are there.

Investigating the bio-assessment and synthesis of compounds with anticancer and DGAT1inhibiting properties is an important and rapidly evolving field at the crossroads of organic chemistry and medicine. In lipid metabolism, the diacylglycerol acyltransferase 1 (DGAT1) enzyme plays a role, specifically in the last stage of triglyceride production. Drug development has taken an interest in DGAT1 because of its possible therapeutic intervention target status, which has arisen due to its involvement in lipid homeostasis. At the same time, finding new cancer treatments is a major problem all around the world; this calls for creative new chemicals that work better and have less side effects. This in-depth analysis of the design and synthesis of DGAT1-inhibiting and anticancer drugs, as well as their biological activity, is presented in a critical review.

To synthesize drugs that inhibit DGAT1, one must carefully plan the chemical structures that will bind to the enzyme and alter its activity. A thorough familiarity with the structureactivity relationship (SAR) is necessary for this procedure in order to maximize the compounds' pharmacological characteristics. Synthetic tactics used by researchers might vary from more conventional approaches like organic synthesis to more novel ones like click chemistry and bio conjugation strategies. In order to find the most effective and scalable



ways to produce these chemicals on a wide scale, it is essential to critically examine various synthetic methodologies.

Concurrently, due to the multidimensional character of cancer biology, the synthesis of anticancer chemicals utilizes a multipronged strategy. To develop chemicals that selectively attack cancer cells while causing minimal harm to healthy cells, structure-based drug discovery, combinatorial chemistry, and rational drug design are crucial. In order to develop effective anticancer drug candidates, it is necessary to strike a balance between the medication's potency and selectivity; this can be difficult to do, as these synthetic techniques demonstrate.

To evaluate their pharmacological characteristics, bio-assessment of DGAT1-inhibiting and anticancer drugs involves a range of in vitro and in vivo experiments. To evaluate the drugs' capacity to alter lipid metabolism, researchers utilize cellular models and enzymatic tests in the setting of DGAT1 inhibition. Given the intricacy of biological systems and the possibility of off-target effects, it is crucial to critically examine these bio-assessment approaches in order to comprehend the validity and applicability of the acquired data.

On the flip side, anticancer drug evaluations include checking for cytotoxicity, apoptosisinducing capabilities, and cancer cell selectivity. To better understand the drugs' effects and possible side effects, preclinical research using xenograft and syngeneic animal models is essential. In order to overcome obstacles such tumor heterogeneity and medication resistance, it is crucial to critically examine these studies and determine whether the results can be applied to clinical settings.

ANABOLISM

Anabolism is a metabolic pathway that builds larger molecules from their constituent parts. Energy storage is maintained by the process, which is associated with the development and maturation of cells, organs, and tissues in the body. Growing new muscles, organs, and tissues is what the anabolic process is all about. The conversion of adenosine triphosphate (ATP) to adenosine diphosphate (ADP) is the process that provides the energy needed for constructive metabolism.

The intricate web of biochemical reactions that make up anabolism—the constructive phase of metabolism—is responsible for the synthesis and construction of molecules essential to the



development, upkeep, and repair of all living things. Contrasted with catabolism, the degradative phase, this complex network of routes and reactions is crucial for life since it allows for the synthesis of vital bio molecules.

The production of large molecules—including proteins, nucleic acids, lipids, and complex carbs—is central to anabolism. The process necessitates the application of energy in order to transcend thermodynamic limits, and it entails transforming simpler molecules into more intricate structures. In order to keep things in check, anabolism is subject to strict regulation by a network of enzymes, hormones, and cellular communication channels.

The complex tango of transcription and translation is the basis of anabolic activities, which include protein synthesis. The genetic code is transported to the ribosomes in the cytoplasm via messenger RNA (mRNA), which is produced in the nucleus by transcription of DNA. Polypeptide chains, which are the building blocks of proteins, are formed when amino acids are linked in a sequence specified by messenger RNA. The intricate nature of anabolic pathways is demonstrated by the level of accuracy and coordination required for this activity.

Another critical component of anabolism is the synthesis of nucleic acids, which include RNA and DNA. DNA replication and RNA synthesis work hand in hand to guarantee accurate genetic information transmission during cell division and the subsequent production of functional proteins. This highlights the significance of anabolism in preserving genetic material, as these activities necessitate careful management to avoid mistakes in the genetic code.

CATABOLISM

In order to release energy, catabolism breaks down big molecules like polysaccharides into smaller ones, called monosaccharides. The breakdown of polymers releases smaller units, which cells can utilize to either build new polymer molecules or degrade them further into waste, generating energy in the process. Thus, the chemical energy required for cellular upkeep and growth is supplied via catabolism. A metabolic reaction is a complex chemical process. A healthy metabolism is defined as the production of an adequate quantity of enzymes to transform food into energy. A metabolic disease can develop when an enzyme is either underactive or overproduced by the organism.



Complex molecules are broken down into simpler pieces during catabolism, the destructive phase of metabolism. This sets off a series of basic and elaborate biochemical reactions that provide energy that is vital for different cellular activities. A steady supply of energy and precursor molecules required to sustain life is maintained by catabolism, the antithesis of anabolism. Central to cellular homeostasis, energy demand response, and recycling of critical building blocks is catabolism, which involves the decomposition of macromolecules like proteins, lipids, carbs, and nucleic acids.

Protein catabolism, the process by which proteins are broken down into their component amino acids, is an essential part of cellular metabolism. Recycling amino acids and removing broken or extraneous proteins are two areas where this mechanism really shines. Proteolysis is the most important process in protein catabolism because it allows for the release of individual amino acids by cleaving peptide bonds. This maintains an endless supply of amino acids, which are necessary for making proteins and other vital macromolecules.

Central to lipid catabolism is the process of lipolysis, which breaks complex lipids into glycerol and fatty acids. Cells rely on lipids as an energy source, and when the need arises, lipolysis lets them access their stored fats. The breakdown of triglycerides into fatty acids can be initiated by hormone-sensitive lipase and other hormonally regulated lipases. Fatty acids can subsequently undergo further metabolism, such as beta-oxidation, to produce ATP.

Catabolism of carbohydrates occurs when complex carbohydrates, especially glucose, are broken down in order to provide energy by means of cellular respiration. The cytoplasm is the site of the first step of glucose catabolism, glycolysis, which produces pyruvate and a trace quantity of ATP. After there, it's on to the mitochondria for pyruvate to go through the oxidative phosphorylation and tricarboxylic acid (TCA) cycle, which results in a much bigger ATP output. Cellular respiration, which includes glycolysis, the TCA cycle, and oxidative phosphorylation, is a key process that cells use to generate energy.

METABOLIC DISORDERS

When a metabolic issue sets in, it's possible that vital nutrients are either over-or underconsumed. As an example of a metabolic illness, diabetes causes excessive blood sugar levels due to either an inadequate insulin response or an inadequate insulin production. When vital organs like the liver, pancreas, and kidneys stop working properly, or when there is a genetic enzyme imbalance, it can lead to metabolic diseases. A malfunctioning metabolism can lead



to an increase in harmful molecules that disrupt normal bodily function or a decrease in the body's capacity to produce necessary compounds. Metabolism disorders can run in families. There is a sizable group of hereditary metabolic diseases known as inborn errors of metabolism. Metabolic mistakes that occur at birth are more commonly known as inherited metabolic diseases or congenital metabolic disorders.

Disruptions to the body's normal metabolic processes, which are responsible for converting food into energy and necessary chemicals, describe a wide range of medical illnesses known as metabolic disorders. Metabolic pathways involved in nutrition breakdown, biomolecule synthesis, and energy production are all susceptible to the onset of these illnesses. Metabolic disorders cause anomalies that can impact various organs and systems; these abnormalities can be inherited, caused by environmental causes, or a mix of the two. Diagnosis, treatment, and continuing research to discover new therapeutic approaches all depend on our ability to fully grasp the complexities of these conditions.

Deficiencies in the body's capacity to absorb and use sugars constitute a notable class of metabolic illnesses known as disorders of glucose metabolism. When insulin synthesis is inadequate or insulin action is impaired, the result is high blood glucose levels, a hallmark of the metabolic condition diabetes mellitus. The immune system's attack on the pancreas's insulin-producing beta cells is the main cause of type 1 diabetes, whereas insulin resistance and insufficient insulin secretion are the main symptoms of type 2 diabetes. Cardiovascular disease, renal failure, neuropathy, and retinopathy are long-term consequences of diabetes that highlight the systemic effect of disturbances in glucose metabolism.

Among the many metabolic illnesses, lipid metabolism disorders stand out. Excessive levels of lipids, especially cholesterol and triglycerides, in the blood, can be seen in hyperlipidemia. Disturbances in lipid metabolism, which are caused by a combination of hereditary factors, food choices, and lifestyle, put people at risk for atherosclerosis and cardiovascular illnesses. An genetic condition known as familial hypercholesterolemia causes decreased clearance of low-density lipoprotein (LDL) cholesterol from the bloodstream due to mutations that impact the LDL receptor. Keeping lipids in a healthy balance is crucial for heart health, as these illnesses show.

TYPE OF METABOLIC DISORDERS



The literature lists a variety of metabolic disorders, and the search for other metabolic disorders is ongoing. Metabolic disorders can be attributed to malfunctions in enzyme or hormone activity, as well as to hereditary abnormalities. But most of these disorders are hereditary, meaning they develop in one family and then spread to another as a result of mutations that cause metabolic abnormalities. Amino acid metabolism disorders, carbohydrates metabolism disorders, organic acid metabolism disorders, etc. are the many types of inherited metabolic abnormalities. Phenylketonuria (PKU), maple syrup urine disease (MSUD), galactosemia, tyrosinemia, homocystinurea, G6PD deficiency, hyperthyroidism, hypothyroidism, peroxisomal disorders, mitochondrial disorders, and many more are among the most prevalent of these.

A wide range of medical illnesses are known as metabolic disorders, and they are all defined by disturbances in the complex network of biochemical reactions that control metabolism. There are several forms of metabolic disorders; these include problems with glucose, lipid, amino acid, and nucleotide metabolism, problems with mitochondrial function, and shortages in particular enzymes. Diagnosis, care, and continuing research into the causes and development of targeted treatments all depend on a thorough understanding of the subtleties of these various metabolic illnesses.

One important group of metabolic illnesses includes abnormalities of carbohydrate metabolism, which includes diabetes mellitus. There are two primary categories of diabetes mellitus: Type 2 diabetes is defined by insulin resistance and insufficient insulin production; Type 1 diabetes is caused by the autoimmune death of insulin-producing beta cells in the pancreas, leading to an insulin deficit. The cardiovascular system, kidneys, nerves, and eyes are all negatively impacted by the high blood glucose levels caused by both forms of diabetes. The increasing number of people diagnosed with diabetes around the world has highlighted the need for holistic methods of care, such as dietary and exercise changes, medication, and, in certain situations, insulin injections.

When lipids, especially cholesterol and triglycerides, are not properly processed and used, it can lead to a variety of illnesses related to lipid metabolism. Elevated blood lipid levels, or hyperlipidemia, are a typical symptom. Disturbances in lipid metabolism, which are caused by a combination of hereditary factors, food choices, and lifestyle, put people at risk for atherosclerosis and cardiovascular illnesses. An genetic condition known as familial hypercholesterolemia causes decreased clearance of low-density lipoprotein (LDL)



cholesterol from the bloodstream due to mutations that impact the LDL receptor. Medications to lower cholesterol levels, changes to one's way of life, and close monitoring of cardiovascular risk factors are all part of the management of problems involving lipid metabolism.

Proteins rely on specific amino acids, which can be either synthesized or broken down in aberrant ways in people with diseases of amino acid metabolism. One well-known disease of amino acid metabolism is phenylketonuria (PKU), which occurs when the enzyme phenylalanine hydroxylase is not present. Intellectual impairments, convulsions, and developmental delays can result from the buildup of phenylalanine caused by this deficit if it is not corrected. Reducing the burden of PKU requires early detection through neonatal screening and dietary therapy, namely decreasing phenylalanine intake. Timely intervention and individualized treatment approaches are crucial for other abnormalities of amino acid metabolism, like homocystinuria and maple syrup urine illness.

Enzymes involved in glycogen metabolism are defective in a class of metabolic illnesses known as glycogen storage disorders (GSDs). Glycogen accumulation or depletion in different tissues is disrupted due to these illnesses, which impacts energy homeostasis. Accumulation of glycogen in lysosomes causes Pompe disease, a form of GSD, which is caused by a shortage of the acid alpha-glucosidase enzyme. Muscle weakness and breathing problems are the main symptoms of this illness. The encouraging results of enzyme replacement therapy in the treatment of Pompe illness highlight the possibilities of personalized medicines for metabolic diseases.

BIO-ASSESSMENT

In order to gain useful insights into the structure, function, and overall health of biological systems, organisms, or substances, bio-assessment is essential in scientific research and medical diagnostics. This comprehensive method makes use of a broad variety of tools, including physiological assessments, behavioral analysis, and genetic and cellular tests. The main objective of bio-assessment is to produce useful information that can support medical diagnosis, decision-making, and scientific investigation. In order to better understand living systems, disease causes, and how to design therapeutic approaches, this thorough investigation of biological entities is crucial.



Analyzing genetic material is a common first step in molecular bio-assessment since it provides a basic grasp of the systems that control cellular functions. Genetic information can be deciphered with the use of molecular tools like polymerase chain reaction (PCR), gene expression profiling, and DNA sequencing. Finding disease-related genes, learning about DNA sequence variants, and investigating how gene expression is regulated under various conditions are all part of this field's purview. To better understand how cells perform and what causes health and illness at the molecular level, molecular bio-assessment is essential.

Proteins, which are essential for cellular function, are also the focus of extensive bio-analysis. The quantification and characterization of proteins are made possible by techniques such as mass spectrometry, enzyme-linked immunosorbent assay (ELISA), and Western blotting. Scientists can learn a lot about cellular signaling networks, new biomarkers, and potential treatment targets by studying protein expression, modification, and interaction alterations. Understanding the mechanisms underlying many diseases, such as cancer, neurological disorders, and metabolic problems, requires molecular-level bio-assessment.

SYNTHESIS AND BIO-ASSESSMENT

In the pursuit of knowledge, comprehension of complex systems, and the development of applications across numerous fields, synthesis and bio-assessment are interdependent aspects of scientific inquiry. The term "synthesis," which has its origins in the chemical sciences, describes the process of making or assembling substances, materials, or structures by means of chemical reactions or other procedures. Bio-assessment, in contrast, is the process of thoroughly examining biological entities, from individual molecules and cells to entire ecosystems, in order to learn more about their make-up, how they work, and how healthy they are. Various disciplines, such as chemistry, biology, medicine, environmental science, and materials science, are involved in this complex relationship between synthesis and bio-assessment, which helps to create novel materials, diagnostic tools, medicines, and sustainable practices.

The synthesis of new compounds with certain desired characteristics is an essential step in the chemical sciences. In organic synthesis specifically, complicated molecules are built up by use of a cascade of chemical processes. The synthesis of organic compounds is fundamental to the fields of medicine, materials science, and cutting-edge technology, and its effects are far-reaching. Organic transformations, catalysis, and multi-step synthesis are just a few of the



synthetic approaches used by chemists to build molecules with the desired properties and functions.

One area where chemical synthesis has had a significant influence on people's well-being is in the production of pharmaceuticals. The field of medicinal chemistry aims to develop and produce compounds that can alter biological functions by focusing on disease-related proteins or pathways. Synthesizing various chemical entities, screening them for biological activity, and improving their pharmacological properties are common steps in the creation of novel medications. An essential part of the pharmaceutical industry's work in finding and developing treatments for various medical issues is the synthesis of small molecules, peptides, and biologics.

An essential part of developing new drugs is doing bio-assessments, which include a wide range of methods for determining how well, safely, and biologically active manufactured chemicals are. To determine the biological impact of synthetic compounds, researchers use in vivo experiments, cellular models, and high-throughput screening tests. The bio-assessment of new compounds reveals important details regarding their medicinal potential, side effects, and action mechanism. Rational drug design is based on this iterative process of bioassessment and synthesis, which helps researchers create safe and effective medicinal interventions.

CONCLUSION

This study aims to shed light on the role of adipose tissue in the development of obesity, a significant metabolic condition characterized by the storage of excess calories as triglycerides. There are currently no medications on the market that can treat obesity other than Orlistat and Lorcaserin. But they do have some negative side effects. To alleviate the impact of these disorders, there is a need for new medications that target different systems and have fewer adverse effects. The last phase of triglyceride (TG) biosynthesis relies on two well-known DGAT enzymes, Acyl CoA: Diacylglycerol acyltransferase-1 (DGAT1) and Acyl CoA: Diacylglycerol acyltransferase-2 (DGAT2). A lack of essential fatty acids causes skin defects in DGAT2 defective mice. As a result, DGAT2 defective mice have a rapid mortality rate after birth. Being able to live depends on this enzyme. Total triglyceride levels were found to be lower in DGAT1 knockout mice, which are otherwise viable. Additionally, when fed a high-fat diet, it showed no signs of diet-induced obesity (DIO). As a result,



treating hypertriglyceridemia and obesity by blocking the DGAT1 enzyme is a promising strategy.

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