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Study of Pharmaceutical Organic Compound: Paracetamol

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ABSTRACT

Paracetamol (acetaminophen) is one of the most widely used pharmaceutical organic compounds used in the treatment of pain and lowering of temperature. Many times the drug is prescribed for medical reasons and concerns as it is inexpensive yet very effective when a patient uses it according to the recommendations. Paracetamol belongs to the chemical class of para-aminophenol and possesses strong analgesic and antipyretic properties. This paper will summarize the information regarding the chemical structure of paracetamol, its physical-chemical properties, synthesis and manufacturing process, the mechanism of action, medicinal usage, metabolism, toxicity, analysis methods, and recent developments. It will pay particular attention to the application of the analytical methods such as HPLC, UV visible spectroscopy, HPTLC, and electrochemical methods of analysis and monitoring. Moreover, the paper will analyze the environmental issues connected with the pharmaceutical waste and present the ways to solve the issue such as green synthesis, nanotechnology, and sustainable pharmaceutical practices. In conclusion, it may be said that the drug in question is an effective medicine which fits the title of the most useful analgesic and antipyretic drug.

Introduction

Medicinal organic compounds are carbon-containing chemical substances which are essential to contemporary medicine because they help to prevent, diagnose, and treat many diseases. They represent the vast majority of medicines utilized in modern pharmaceuticals due to their variety of chemical forms, effects, and despite the reliability of pharmaceuticals. In the past century, important progress in organic chemistry, medicinal chemistry and pharmacy have allowed creating better and safer drugs. Today, medicinal organic compounds are used in the majority of healthcare systems throughout the world. Among the various medicinal organic compounds, paracetamol (acetaminophen) is perhaps the most widely used medication available by prescription and over the counter. It is part of the World Health Organization (WHO) Model List of Essential Medicines due to its high efficacy, low cost and its favorable safety characteristics when taken in appropriate dose. Since the day of its introduction in the 20th century, paracetamol has proven itself to be one of the most widely used medications in the world and billions of doses are administered each year. Paracetamol is an organic substance with the chemical formula of N-(4-hydroxyphenyl)acetamide, whose molecular weight is 151.16 g/mol. It belongs to the family of para-aminophenol derivatives and exhibits advanced analgesic and antipyretic characteristics. Paracetamol is unique among many NSAIDs for its lack of anti-inflammatory properties, however, it is able to efficiently deal with mild-to-moderate pain and treat fever. Moreover, its gastrointestinal side effects may be described as negligible. Thus, this over-the-counter drug is advised for patients who are unable to tolerate other NSAIDs drugs. It seems that paracetamol operates through the inhibition of prostaglandins production in the central nervous system causing a diminished pain response and the regulation of temperature. This drug mechanism is not yet completely understood, but there exist certain hypotheses

claiming that the overall effect of paracetamol is mainly realized by a central cyclooxygenase inhibition. The chemical structure of paracetamol is fairly simple and provides high availability for the needs of pharmaceutical industry. New methods of synthesis to increase the efficiency of paracetamol are being studied all the time. Paracetamol has a good safety record concerning therapeutic doses, but if taken in excess, there are severe possible consequences for human liver.

The objective of the review was to examine paracetamol from various perspectives of its chemistry and synthesis, physicochemical principles and effects, mechanisms and application, methods of identification and measurement of toxicity, environmental issues concerning the application of the substance and its other aspects.

Literature Review

Paracetamol, known internationally by its generic name acetaminophen, is one of the most commonly used pharmaceutical organic compounds in medicine today due to its special analgesic (pain-relieving) and antipyretic (fever-reducing) effects. Paracetamol has been in clinical use since the middle of the twentieth century. Since then, it has become one of the most commonly prescribed and used medications in the world. The use of paracetamol for the treatment of mild to moderate pain and fever is supported by its therapeutic effectiveness, long history and low price. Due to its advantages, paracetamol has been included in the WHO Model List of Essential Medicines as one of the most important drugs for every healthcare system. Paracetamol is a derivative of para-aminophenol. The drug acts differently from many non-steroidal anti-inflammatory drugs (NSAIDs) in that it does not have a damaging effect on the gastrointestinal tract and affect platelet aggregation. Therefore, it is used in children, pregnant women under medical supervision, and people who can't tolerate aspirin and other NSAIDs. Common conditions treated with paracetamol include headache, toothache, muscle pain, joint pain, postoperative pain, menstrual pain, influenza

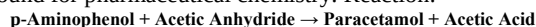
and other febrile diseases. From the pharmaceutical viewpoint, paracetamol has many favorable properties, like the fast absorption in the gastrointestinal tract and great bioavailability. It is more widely used as a drug because the medicine is administered easily in many pharmaceutical forms like tablets, capsules, syrup, suspensions, suppositories, IV injections, and disperse tablet forms. In contrast to the growth of the market demand for safe and effective analgesics, the scientific research of paracetamol is going on. Nowadays it is conducted in many areas, including the improvement of therapeutic efficacy and the enhancement of formulation technology, quality control, analytical determination, controlled drug delivery systems, and eco-friendly manufacturing.

Chemical Structure and Physical and Chemical Properties

Paracetamol, otherwise known as N-(4-hydroxyphenyl)acetamide (or 4-hydroxyacetanilide), has a molecular formula $C_8H_9NO_2$, and its weight is equal to 151.16 g/mol. The moderate solubility of paracetamol is advantageous in terms of its pharmaceutical formulation since it ensures sufficient dissolution and absorption required for the therapeutic action after oral administration.

Paracetamol has a melting point of 169-170 degrees Celsius which shows good thermal stability of the compound during its production and storage. The chemical stability of paracetamol influences by the storage conditions such as temperature, humidity, and light.

The physicochemical properties like stable melting point, moderate solubility, and lipophilicity make paracetamol a good model compound for pharmaceutical chemistry. Reaction:



The process of purification of the product will start immediately after its production, as after synthesis it needs purification and recrystallization to get paracetamol that meets the production standards. After being purified the crystals will be filtered and dried before being milled and analyzed for compliance with the quality standards (GMP). The production of paracetamol in the pharmaceuticals industry is heavily regulated and subject to quality control and regulation. Qualitative testing of the drug includes identification of the substance purity tests and assay. The last years scientists working in the sphere of paracetamol production are focused on greener processes, which means that environmentally friendly methods of production with fewer toxic chemicals are applied, which means sustainable production. Although paracetamol is widely utilized in modern medicine and is believed to be effective painkiller and antipyretic agents, its enzyme is still under question. The most recent studies show that paracetamol acts. Paracetamol produces its analgesic effect primarily by blocking cyclooxygenase (COX) activity in the brain, leading to diminished awareness of pain. In addition, studies indicate that its analgesic activity could be due to the impact of serotonergic systems or the involvement of metabolite AM404 in modulation of pain transmission. Paracetamol causes its antipyretic effect by acting on hypothalamic thermoregulation center, causing heat dissipation and decrease in body temperature during fever. Paracetamol has a relatively weak anti-inflammatory effect as compared to NSAIDs owing to its inability to exert its COX inhibiting effect because of the presence of high levels of peroxides in the inflamed tissues. This explains the usefulness of paracetamol in relieving pain and fever yet its limited value in cases of severe inflammatory disorders. Paracetamol is rapidly absorbed after oral administration, metabolized in the liver, and cleared from the organism through the kidneys. It is safe when used at recommended dosing, however, misuse can lead to the production of toxic metabolite N-acetyl-p-benzoquinone imine (NAPQI) that may cause irreversible damage to the liver.

Pharmacological Applications- Paracetamol is one of the most widely prescribed medications for pain and fever treatment. It is often used for headache, toothache, muscles ache, back pain, osteoarthritis, postoperative pain, menstrual pain, flu, common cold, and different febrile diseases. With its favorable safety profile and low risk of gastrointestinal side effects, paracetamol is Paracetamol is available in a variety of forms such as pills, injections, and syrups. All these forms enable proper administration of medication based on a person's age and health. In the hospital, injectable paracetamol is a necessary option for the immediate relief of pain when other methods of taking medication are impossible.

Paracetamol is also often part of some certain mixtures, which improves the efficacy of the pain treatment. It must be taken into consideration that misuse of paracetamol may lead to serious

overdose cases, which are one of the most important reasons for drug-induced liver damage.

Assessing Paracetamol using Analytical Methods- Satisfactory analytical methods must exist for the quality assurance and control of medicines containing this active ingredient. There are plenty of methods that can be applied for the identification of the active component. Various methods exist, but HPLC is considered the method of choice due to its unprecedented specificity and accuracy. Moreover, as for the significant techniques, one can speak about High-Performance Thin-Layer Chromatography (HPTLC), Gas Chromatography–Mass Spectrometry (GC–MS), Fourier Transform Infrared Spectroscopy (FTIR), and electrochemical sensors. Each of these techniques has its own advantage for various analytical purposes. Recent progress in analytical chemistry has led to the creation of a new generation of biosensors and electrochemical biosensors made of nanomaterials that make it possible to detect very small concentrations of paracetamol in pharmaceutical products, human fluids, and environmental samples. This has significantly improved pharmaceutical quality evaluation and clinical investigation.

Cutting-Edge Research Directions- Current pharmaceutical research is aimed at enhancing the therapeutic efficacy and safety of paracetamol through innovative formulations. The provided system of drug administration suggests the emergence of several means, such as nanotechnology-based delivery systems, sustained-release tablets, orally disintegrating tablets, or controlled-release formulations.

Green chemistry has also gained popularity. Innovative, eco-friendly synthetic methods aimed at minimizing solvent-based processes, hazardous chemicals, and industrial waste are being developed now. Moreover, computer technologies, including molecular modeling and artificial intelligence, are used to create more efficient formulations. Modern analytical methods made it possible to simplify paracetamol residue detection in wastewater and the environment, which enables us to investigate pharmaceutical pollution in our environments.

Research Gaps- Although paracetamol has been comprehensively researched over decades, some important obstacles still exist. The molecular mechanism that is responsible for its pain-relieving action has still to be investigated, as well as the pharmacological pathways of the drug. Another aspect requiring careful investigation is the safety of the drug during long-term use, especially among patients suffering from liver insufficiency or other liver diseases.

The rising level of pharmaceutical contamination of water bodies has raised new concerns about the environmental persistence of paracetamol and its products of degradation. Hence, it has become clear that more focus must be given to the sustainability of production, wastewater treatment, and environmentally safe disposal methods. A lot of attention should also be directed at creating greener synthetic processes, highly selective analytical methods, and advanced systems of drug administration. In conclusion, the development of modern technologies, involving pharmaceutical chemistry, medicinal chemistry, analytical chemistry, nanotechnology, and ecology as a whole, will improve the safety and effectiveness of paracetamol.

Methodology

Research Design- The research mentioned was conducted as a systematic review of existing knowledge on the scientific significance of paracetamol as a pharmaceutical organic chemical. Unlike experimental studies, this investigation did not make use of laboratory synthesis or live trials or animal studies, but depended entirely on a systematic process of compiling and evaluating previously published scientific literature. Qualitative research design was employed for the purpose of having the ability to collect information from various scientific studies and having an extensive understanding of the subject. The procured literature was critically analyzed with the aim of revealing the latest scientific advancements, technological developments, and opportunities for future research on the given topic. Information collected from different sources was compared and synthesized in order to provide the evidence-based review that would show the current state of research on paracetamol. The method outlined above also made it possible to reveal similarities, differences, and gaps in the investigation of the published literature. The review was organized in a systematic way to ensure the continuity of information in the different sections of the study. Each topic was analyzed in order to keep a logical sequence from the simplest that is the chemistry of paracetamol to its pharmaceutical use, then to

analytical determination and safety concern, and finally the scientific innovations.

Sources of Scientific Literature-The sources of scientific literature that are used in this review are sourced from peer-reviewed journal articles and reputable academic databases. To ensure the quality of the review many reputable sources have been used. Those sources include Google Scholar, PubMed, Science Direct, Springer Link, Wiley Online Library, Taylor&Francis Online, MDPI Journals, Research Gate, and some other resources. Many authoritative pharmaceutical textbooks and actual reports published by various organizations such as WHO (World Health Organization) were checked to provide and check needed information. The emphasis was focused on peer-reviewed research articles because they have gone through several stages of review, and the latest review articles were considered to see the trends of current changes. However, some important historical publications mentioning the discovery and the technique of preparation of paracetamol should be included if they contain some noteworthy information on its history or something that is connected to the topic of research. To keep the review relevant, a priority was given to the last fifteen years' publications, however, some veteran works that played an important role in the study of paracetamol as well as paracetamol chemistry should have been previously used. Hence a balanced approach to choosing the sources was applied in the course of this work.

The sources of literature are found through a systematic literature search based on the key terms and search phrases. Some of the significant keywords utilized in this study are paracetamol, acetaminophen, pharmaceutical organic compounds, medicinal chemistry, drug synthesis, pharmaceutical analysis, analgesic drugs, antipyretic drugs, metabolism of drugs, toxicity, HPLC, UV-Visible spectrophotometer, electrochemical analysis of drugs, pharmaceutical formulation, and quality control. Boolean operators like AND, OR, and NOT were utilized in the search. Examples of the combinations used to yield relevant publications include "paracetamol AND pharmaceutical analysis," "acetaminophen AND medicinal chemistry," and "paracetamol AND analytical methods." The duplicate entries retrieved from different databases were eliminated based on careful analysis prior to performing the detailed assessments. The preliminary screening of publications was performed based on their title and abstract. The articles that appeared to be relevant to the objective of this study have been evaluated in detail in order to understand their significance in the review article. Overall, the systematic approach to information retrieval helped to achieve overall comprehensiveness and validity of the information that was used in this review article. Once the literature search was completed, all retrieved materials were screened according to already established inclusion and exclusion criteria. It was performed in order to ensure that only the valid scientific papers contributed to the current review.

- Peer-reviewed research papers published in reputable international journals.
- Review articles covering the chemistry, pharmacology, analytical techniques, and pharmaceutical applications of paracetamol.
- Papers published in the English language.
- Reports containing valid experimental data and scientifically proven conclusions.
- Standard pharmaceutical reference books and official guidelines on paracetamol.
- Articles that are indexed in different data sources.
- Paper presentations of conferences that do not contain final results.
- Reports, press notes and editorials.
- Investigations irrelevant to paracetamol and various medicines.
- Papers with inadequate scientific evidence and imperfect methodologies.

Prior to the organization of the information, the necessary papers were classified according to their themes that relate to the investigations and will provide the necessary information on chemical properties of the drug, the chemical synthesis, the way the substance operates, the analytical procedures, pharmacological orientation of paracetamol, toxicity, social consequences and new scientific achievements that have taken place in the given field.

Data Extraction and Systematization After the literature review, the process of systematic data extraction was initiated, including all the data obtained in the research. The process of data extraction concentrated on such aspects as the chemical structure, the basic

physicochemical characteristics of the drug, ways of obtaining it, the mechanism of paracetamol, its effects in medicine, and methods of analysis. In order to achieve high quality during the review, collected facts were systematized according to a given theme. The similarities across the literature on paracetamol were juxtaposed and contradictory info analyzed and discussed this strategy has increased the accuracy and trustworthiness of the review while keeping in check that the discussion concentrated on the chosen research subject.

Comparative Study of Previous Research

A comparative study was conducted to analyze the similarities, discrepancies, benefits, and shortcomings revealed by earlier scientific research. Different studies were compared regarding their aims, research methods, techniques, activity, and conclusions. Special attention was paid to the comparison of various methods of analysis to determine paracetamol such as High-performance liquid chromatography (HPLC), UV-visible spectrophotometry, high-performance thin layer chromatography (HPTLC), gas chromatography-mass spectrometry (GC-MS), and other electrochemical methods of analysis. Their sensitivity, accuracy, precision, detection limit, advantages, and applicability have been critically assessed. Besides, the studies were compared with respect to the pharmaceutical formulations used in research, synthesis methods, and delivery systems. Such a comparison provided an opportunity to determine the most effective methods available today and to detect the areas that need further searching.

Quality Evaluation of Selected Literature-In order to make sure about the reliability of scientific publications, each of the publications that was included in the structure of research was rigorously evaluated. The first preference was given to articles published in scientific peer-reviewed magazines since they are very strictly checked before being published. More importance was attached to the study where the goals of the research were precisely provided. The trustworthy scientific research was based on trustworthy experimental data backed by certain pharmaceutical standards. Reputable journals' reviews were examined thoroughly as well since their articles include results of independent scientific studies but summarize valuable information on modern scientific developments. Thus, the process of assessing the quality of the literature has added prominence to the review.

Data Analysis-The information collected and analyzed was carefully interpreted based on evidence. The results situated in different publications were analyzed to obtain the main scientific trends and advancements in the problem of paracetamol research. The analysis took into account the correlation between the chemical property, physicochemical properties, methods of synthesis, pharmacological activities, and the results of analysis.

The results have been analyzed objectively without any subjective opinions.

Limitations of the Research-The given review provides a summary about paracetamol research as a very important organic compound. Various chemical issues, such as chemistry, synthesis, physicochemical properties, pharmacological characteristics, analysis, toxicity, and environmental issues were discussed. However, the review has some limitations. It does not include the original laboratory experiments performed by the writer and relies only on secondary sources. The results depend on the quality of older published works of scientists. Besides, only publications written in English were taken into consideration. Nevertheless, this review remains credible, scientific, and useful for academic and practical purposes thanks to the careful selection of available scientific magazines, peer-reviewed journal articles, and well-known scientific literature. The given methodology helped to understand the pharmaceutical importance of paracetamol and made it possible to outline some directions for further research.

Results and Discussion

Comparative Evaluation of Paracetamol as an Organic Pharmaceutical Compound-Based on the argumentative analysis of the scientific literature, it can be concluded that paracetamol continues to be among the essential organic pharmaceutical compounds utilized in modern medicine. The comparative evaluation of the previous findings shows the drug has a good combination of therapeutic value, safety, cost, and availability, which is why this medicine is widely accepted as an effective anodyne and antipyretic medicine throughout the world. Compared to many of the traditional pain-treated medications, paracetamol is an efficient cure for mild and

moderate pain with much lower occurrences of other side effects of medication taken or the medicine having fewer negative effects on the gastrointestinal tract. The review of the available academic sources shows that due to its simple molecular structure paracetamol can be synthesized easily and made available in various dosage forms across the medical arena. Among the most common forms of this medicine are tablets, capsules, suspensions, syrup, suppository, and injection form. The analysis indicates that the process of synthesis of paracetamol is effective due to its good stability in terms of chemicals. The literature indicates that its effectiveness in the given form depends directly on its proper administration and the administering process itself. It is determined that the appropriate dosage results in effective relief in pain and fever; however, its prolonged or excessive intake increases. Based on the overall analysis, it can be concluded that paracetamol is still in demand for its extensive array of therapeutic uses, good safety profile, and proof of efficacy in clinical practice. As indicated in the analysis of the relevant literature, there is a need for accurate measurement of paracetamol for the purpose of quality control in the pharmaceutical industry, clinical applications, and ensuring regulatory compliance. The results of the analysis of various analytical methods indicate that High-Performance Liquid Chromatography (HPLC) is the most reliable technique. HPLC is very widely used in the pharmaceutical industry for assay determination, impurity profiling, stability testing, and quality control. One more technique called UV-visible spectrophotometry is simple, fast, and cost-effective; however, its sensitivity is lower than that of HPLC. The high sensitivity of HPLC makes it possible to obtain reliable and reproducible results even in education institutions as well as for the routine quality control applications.

Meanwhile, there are modern methods like HPTLC, GC-MS, and electrochemical sensors that are effective for measuring paracetamol. Electrochemical techniques are accepted widely due to their high speed of analysis and low costs. There are recent researches on the applications of nanostructure-based electrochemical sensors for detecting low levels of paracetamol in pharmaceuticals and environment.

Pharmacological Activity and Therapeutic Importance- It is obvious from the studies analyzed that paracetamol continues to be one of the best non-opioid analgesics and antipyretics. Due to the fast acting mechanism of action, good level of tolerability, and low incidence of adverse effects in the digestive tract, paracetamol is often the best choice for treating mild pain or fever. Clinical data show that paracetamol in therapeutic doses takes away headache, toothache, musculoskeletal pain, post-operative pain, symptoms of influenza, and fever in adults and children.

Paracetamol is safer than many non-steroidal anti-inflammatory medications, because it is virtually non-irritating to the stomach. All this helps elderly, young patients, and those who cannot take NSAIDs. It can be seen from the literature studied that paracetamol is taken with other analgesics to improve pain management.

Despite the benefits of paracetamol.

New Progress in Pharmaceutical Science

The latest studies in pharmaceutical science greatly assisted in improving paracetamol effectiveness in healthcare. At present time, the primary pharmaceutical development area is related to drug stability; bioavailability; and better patient adherence facilitated by the inventions of new drug formulations such as sustained-release tablets; orally dissolving drugs; pediatric formulations and intravenous administration.

Nanotechnology became a very trendy subject in pharmaceutical science with many delivery systems being developed. The drug delivery techniques utilizing nanoparticles and new-generation polymers are expected to facilitate controlled drug release; the efficacy and the reduction of side effects.

The development of accurate analysis methods using nanomaterials and electrochemical techniques greatly contributed to pharmacological manufacturing as paracetamol can be easily detected in medicines and biological substances.

Environmental and Pharmaceutical Issues- Due to the increasing popularity of paracetamol, its presence in the environment becomes quite worrying. The information about paracetamol being found in sewage, rivers, lakes, and hospital wastes has been published. The concentrations of the drug are not very great but the repeated emission of pharmaceutical products into the water bodies can affect

microorganisms and aquatic animals. Also, it is highly important to get rid of expired medications properly. The wrong disposal contributes to pollution and contamination of the environment, that is why scientists recommend using green technologies in the production; wastewater treatment processes; and methods of disposal of pharmaceutical waste for this problem.

Another problem in pharmaceutical production is the necessity to maintain the same quality of products. All authorities have created laws and regulations concerning making sure of compliance with GMP during manufacturing.

Future Expectations- According to scientific literature, the further studies towards paracetamol should be devoted to increasing its pharmaceutical efficiency and improving the environmental situation. The green synthesis technologies; research methods and drug delivery methods should be developed and implemented on the market.

The application of artificial intelligence and computational chemistry is predicted to speed up the developments in accordance with the drug formulation; stability predictions; and optimization processes. Nanotechnologies are expected to increase the therapeutic power and make it possible to minimize side effects of drugs.

In general, the facts indicate that paracetamol will preserve its significance for medicine in the nearest future along with improvements and progress in the pharmaceutical sphere.

Conclusion

Paracetamol is among the active organic substances widely used as a medicinal compound. This review suggests that paracetamol is one of the most used drugs due to its analgesic and antipyretic effects and indicates that it has several advantages that make it suitable for medicine (**Elias et al., 2020**). The review also indicates the impact of pharmaceutical chemistry on the study of paracetamol. Thus, the development of pharmaceutical chemistry has led to a considerable increase in knowledge about paracetamol in terms of its synthesis, mechanism of action, physicochemical properties, formulation, and applications. Paracetamol has many different advantages over other drugs, such as simple chemical structure, good physicochemical properties, high therapeutic activity, patient acceptability, and safety when used in the recommended amount. At the same time, it should be noted that paracetamol is very safe when used in therapeutic doses; at the same time, the abuse and overdose of this drug is quite dangerous, since it can lead to the development of liver toxicity. To sum up, it can be stated that rational prescriptions, patients' awareness about prescribed doses, and systematic pharmacovigilance are necessary for minimizing the number of adverse reactions occurred due to wrong treatment and making clinical application of any drugs safe and efficient. The pharmaceutical industry and healthcare professionals have to pay more attention to ensuring quality of pharmaceuticals, compliance with the regulations, and informing the patients about their medication. The important conclusion that can be done based on the reviewed literature is the increasing concern about pharmaceuticals pollution of the environment. Paracetamol is widely consumed nowadays, which in turn results in its occurrence in sewerage and aquatic ecosystems. Thus, it is highly critical to implement green technologies in production and disposal of drugs. Modern discoveries such as those related to nanotechnology, controlled drug delivery systems, computational drug design, artificial intelligence, etc. have opened new doors for improving paracetamol's therapeutic effect and safety. Said technologies will help improve stability of medicines, their bioavailability, and introduce new pharmaceutical technologies. Finally, the findings of the review show that paracetamol will undoubtedly remain one of the most relevant pharmaceutical organic compounds because new research is needed to invent safer types of drugs and ecologically friendly technologies, sensitive analysis, and drug delivery systems.

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