

The Development of Gold Nanomaterials for Cancer Treatment and Diagnosis

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Abstract. Nowadays, gold - based nanomaterials have emerged as a highly promising candidate for cancer treatment, garnering significant attention. Their distinctive optical, electronic, and physicochemical characteristics endow them with diverse therapeutic approaches. For instance, photothermal therapy mechanisms, gold nanomaterials are capable of efficiently converting light energy into heat. Under laser irradiation, they can selectively eliminate cancer cells. Moreover, these nanomaterials can serve as drug carriers for targeted drug delivery. By doing so, they enhance the effectiveness of chemotherapy drugs while reducing the side - effects associated with them. This paper provides an overview of the most recent breakthroughs in cancer treatment using gold nanomaterials.

Keywords: Cancer; Gold Nanomaterials; Treatment; Diagnosis.

1. Introduction

Cancer, a truly menacing term, once it intrudes into people's lives, triggers waves of upheaval and callously alters the fates of countless families. Cancer cells in the human body grow without being governed by normal physiological mechanisms, exhibiting the characteristic of unrestrained proliferation. They can not only keep growing and form tumors at the primary site but also invade the surrounding normal tissues and spread throughout the body via the circulatory or lymphatic systems. This process, known as metastasis, is the main cause of cancer - related fatalities.

Despite cancer being a major disease that severely threatens human health, with the continuous progress of medical science and the gradual deepening of people's understanding of it, remarkable advancements have been made in cancer prevention, diagnosis, and treatment. Through effective preventive measures, early - stage screening, and standardized treatment, an increasing number of cancer patients can achieve better treatment results and quality of life. Scientific materials play a crucial and ever - more - important role in this context, being relevant to food, clothing, and daily life. Beyond standard process technology and equipment innovation, the development of futuristic new materials often brings about significant progress in the high - tech industry.

Many materials can be modified at the microscopic level. Nanomaterials, for example, refer to materials where at least one dimension in three - dimensional space is at the nanometer scale (1 - 100 nanometers) or is composed of such basic units. This scale corresponds to approximately 10 to 1000 atoms closely packed together. Nanomaterials possess special physical and chemical properties and are classified into metal nanomaterials, inorganic nanomaterials, and organic nanomaterials. Gold - based nanomaterials contain nanoparticles of various metal elements like gold, silver, and copper. These materials have excellent electrical conductivity and good chemical stability, and are widely applied in the electronic and catalytic fields.

Among them, gold nanomaterials play an important function in curing cancer, drug delivery, photothermal effect and so on. For instance, gold nanoparticles are effective drug delivery carriers, enabling precise and regulated release of therapeutic substances. This capability allows for the reduction of adverse effects while optimizing the effectiveness of treatment. In addition, photothermal therapy utilizes the distinctive capability of cancer cells to convert light into localized heat, resulting in the targeted destruction of these cells while minimizing damage to surrounding healthy tissue.

The development of gold nanomaterials has a long - standing history. The research on nano - metal materials can be traced back to the early 20th century. The earliest study on them dates to 1917 when the German scientist Richard Zsigmondy first prepared nano - sized gold particles. In the following

decades, with the development of modern analytical techniques such as electron microscopy and X-ray diffraction, the study of nano-metal materials advanced further. Since the 1980s, with the emergence of nanotechnology, the research on nano-metal materials has entered a phase of rapid development. Researchers have discovered that nano-metal materials have properties and characteristics distinct from traditional metal materials, such as the size effect, surface effect, and quantum effect. These novel properties open up new application possibilities for nano-metal materials, such as in optoelectronic devices, sensors, and catalysis. There are multiple ways to prepare gold nanoparticles, mainly including the sodium citrate reduction method and the electrochemical method.

In 1951, Turkevich used citric acid to reduce chlorauric acid for the synthesis of gold nanoparticles, and this remains one of the commonly used methods. The electrochemical method uses a gold plate as the anode, a sacrificial anode electrode to generate gold ions through electrolysis, and a platinum plate as the cathode to reduce these ions. This method can control the size of gold nanoparticles, but it is energy-consuming.

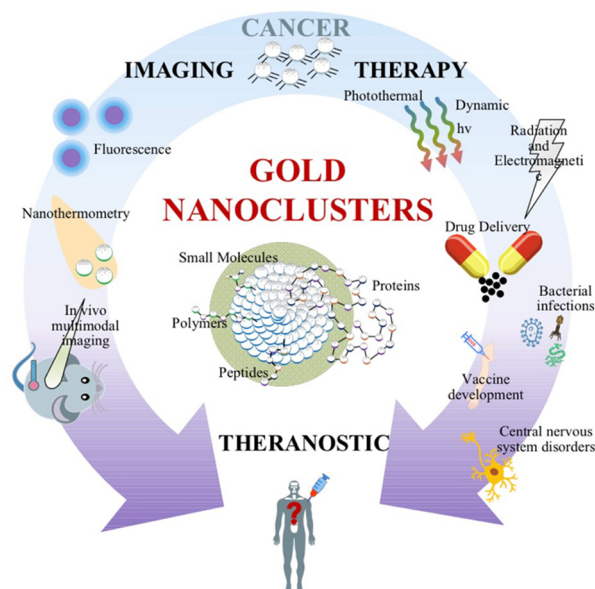


Figure 1. nano-metal materials have shown strong potential and broad application prospects in many fields

Although nano-metal materials have shown strong potential and broad application prospects in many fields, they still face some challenges and problems in practical applications. For example, the preparation method and cost of nano-metal materials are one of the difficult problems to be solved in the current research. At present, most of the preparation methods of nano-metal materials are still complicated, time-consuming and resource-consuming, which leads to high preparation cost and limits their wide application in industrial production. Secondly, the impact of gold nanomaterials on the environment should be considered. Because gold nanomaterials are toxic, they are harmful to organisms and the environment. Environmental considerations should be further strengthened in the use and production of gold nanomaterials, so biocompatibility is also one of the factors to be considered. There are many challenges in the research and preparation of gold nanomaterials. Continuous and deep innovation is needed to solve it. Only by overcoming these challenges can gold nanomaterials be better used in various fields and make greater contributions to the progress of science and technology. In the future, nano-metal materials will not only have outstanding mechanical properties but also possess multiple functions, such as magnetism, optics, and biomedicine. By adjusting their structure and composition, multi-functional design and preparation can be achieved, expanding their application areas. With the growing emphasis on sustainable development, the future preparation of gold nanomaterials will adopt more environmentally friendly and low-energy-consumption technologies to reduce the environmental impact. Most importantly,

the development of nano - metal materials has promoted the research and application of new materials, provided essential support for future scientific and technological development, and advanced the progress and development in the field of materials science and engineering.

2. The Synthesis of These Materials Can be Achieved Through Various Methods, Each with its Own Set of Advantages and Characteristics.

A frequently - employed method is the chemical reduction approach. In this method, gold ions are reduced in the presence of a stabilizing agent to create nanoparticles. This technique enables accurate control over the size and shape of the nanoparticles, which are key elements in their application to cancer treatment. For example, smaller nanoparticles, commonly known as nanoclusters, show distinct properties when compared to larger particles. The size effect is of great significance in cancer therapy as it affects the biodistribution, cellular uptake, and the ability of the nanoparticles to penetrate tumor tissues. Smaller nanoparticles might have superior penetration capabilities, yet they could be removed from the body more quickly. Conversely, larger particles may have longer circulation times but face limitations in penetration. Another important aspect of the synthesis process is the control of the nanoparticles' shape. Various shapes, including nanospheres, nanorods, nanostars, and nanoflowers, have been investigated. Nanospheres are the most straightforward form and have been extensively studied because of their easy synthesis and potential for surface modification. Nanorods, however, have unique optical properties due to their non - isotropic shape. This makes them valuable in photothermal therapy, as they can convert light energy into heat to eliminate cancer cells. Nanostars and nanoflowers provide an increased surface area. This can be advantageous for loading drugs or attaching targeting ligands, thus enabling more efficient delivery to cancer cells.

1) The shift from gold nanoclusters to larger particles is a captivating area of research in cancer therapy, mainly because of the prominent size - related effects. Gold nanoclusters, usually made up of several to several dozen gold atoms, have unique characteristics that set them apart from larger particles. Their small size facilitates enhanced cellular uptake via endocytosis, as they can more readily traverse the cell membrane and intracellular pathways. This feature makes them appealing for the targeted delivery of therapeutic substances directly into cancer cells. Furthermore, the size of gold nanoparticles affects their interaction with biological molecules and cells. Smaller nanoclusters might have a higher surface - to - volume ratio, which could result in increased reactivity and potential toxicity. Conversely, larger particles may have more stable surface chemistry, reducing the likelihood of unwanted interactions. In cancer treatment, the size effect also influences the efficiency of photothermal therapy. Smaller nanoparticles may absorb light less effectively, leading to lower heat production, while larger particles can absorb more light and generate enough heat to trigger apoptosis in cancer cells. Thus, understanding and optimizing the size of gold nanomaterials is essential for customizing their properties for specific cancer therapy uses, balancing aspects like tumor penetration, cellular uptake, circulation time, and therapeutic effectiveness.

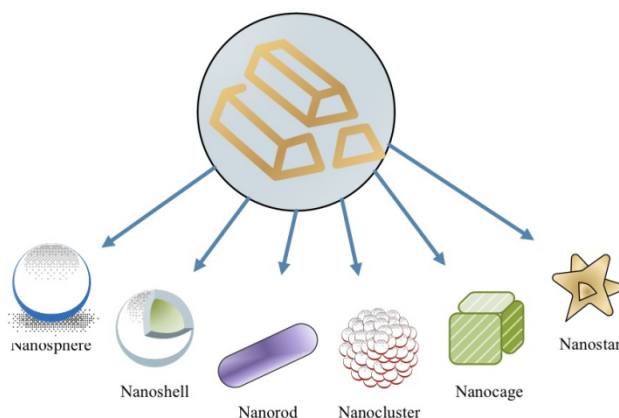


Figure 2. gold ions are reduced in the presence of a stabilizing agent to create nanoparticles

2) The shape of gold nanoparticles has a far - reaching impact on their application in cancer therapy, as different shapes bestow distinct properties and functions. Nanospheres, the most common and simplest shape, provide a large surface area for functionalization with targeting ligands, drugs, or imaging agents. This makes them well - suited for targeted drug delivery, where surface modifications can boost the nanoparticles' specificity towards cancer cells. For instance, attaching antibodies specific to cancer cell surface markers can enable the nanospheres to selectively bind to and be taken up by cancer cells, delivering the payload precisely where it is required. Nanorods, with their elongated form, display unique optical properties due to the plasmon resonance effect. This characteristic makes them highly effective in photothermal therapy. When exposed to near - infrared light, which can penetrate deeper into tissues than other wavelengths, nanorods can absorb the light and convert it into heat with high efficiency. The heat generated can selectively destroy cancer cells while minimizing harm to surrounding healthy tissues. Moreover, the aspect ratio of nanorods can be adjusted to optimize their optical absorption properties, further enhancing their therapeutic potential.

Nanostars, characterized by their branched structure, offer an even larger surface area compared to nanospheres and nanorods. This increased surface area provides more sites for attaching multiple functional components, allowing for the combination of different therapeutic methods on a single nanoparticle.

Nanoflowers, with their complex and irregular structure, have shown potential in cancer biomedical diagnosis. Their unique morphology can lead to enhanced light scattering, making them useful in optical imaging techniques for detecting cancer cells or tumors. The rough surface of nanoflowers can also offer better anchoring points for biomolecules, improving the stability and specificity of nanoparticle - based diagnostic assays.

3. The Functionalization of Gold Nanomaterials

The functionalization of gold nanomaterials is a vital step in fully harnessing their potential for cancer treatment and diagnosis. When specific functional groups or molecules are attached to the surface of gold nanoparticles, their properties can be precisely adjusted to meet the needs of various cancer research applications.

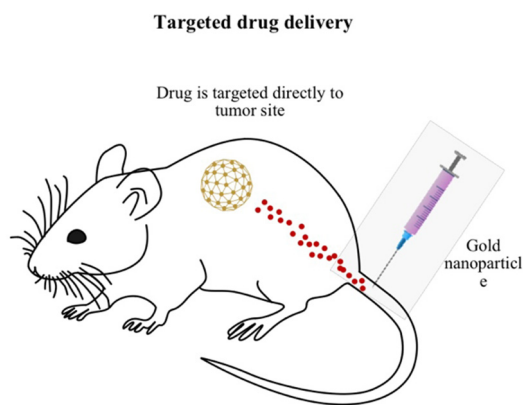


Figure 3. targeted drug delivery

1) One of the primary applications of functionalized gold nanomaterials in cancer therapy is targeted drug delivery. Ligands like antibodies, peptides, or small molecules that can specifically bind to cancer cell receptors can be linked to the nanoparticles. This targeted method ensures that the drugs loaded onto the nanoparticles are selectively delivered to cancer cells, reducing the side effects on normal tissues. The functionalization of gold nanomaterials for targeted cancer treatment not only enhances the therapeutic index but also paves the way for personalized medicine. By identifying the specific biomarkers present in a patient's tumor, nanoparticles can be customized with the right

targeting ligands to offer individualized treatment plans. This approach maximizes the likelihood of successful cancer therapy while minimizing negative impacts.

Functionalization of gold nanomaterials represents a crucial step in unlocking their full potential for cancer therapy and diagnosis. By attaching specific functional groups or molecules to the surface of gold nanoparticles, their properties can be precisely tuned to meet the requirements of different cancer research applications.

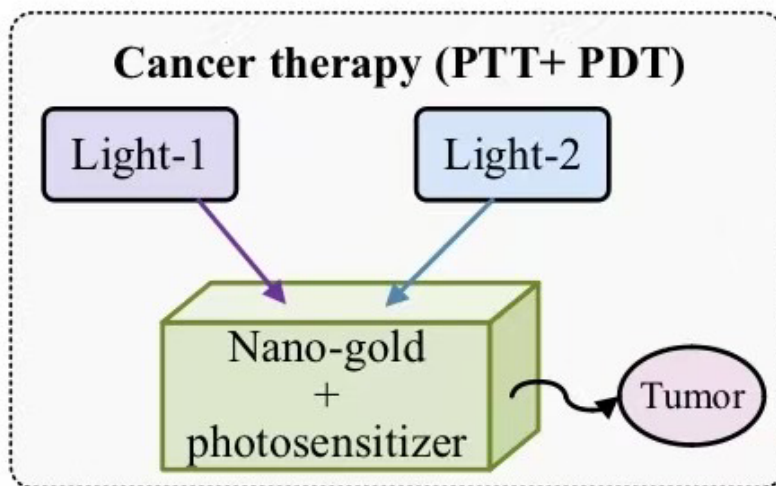


Figure 4. The combination of functionalized gold nanomaterials in photothermal and photodynamic cancer therapies

2) Functionalized gold nanomaterials have come to the forefront as potent tools in photothermal and photodynamic cancer therapies. These materials present innovative ways to treat cancer, boasting high specificity and minimal invasiveness.

In photothermal therapy (PTT), functionalized gold nanomaterials, especially gold nanorods and nanostars, are of utmost importance because of their remarkable light absorption capabilities in the near - infrared (NIR) region. The NIR light has the advantage of being able to penetrate deeper into tissues compared to other wavelengths, making it perfect for targeting tumors located deep within the body. When functionalized gold nanoparticles are exposed to NIR light, they transform the absorbed light energy into heat. This causes a localized temperature rise in the tumor area. The increase in temperature can inflict irreversible damage on cancer cells, ultimately leading to cell death through processes like protein denaturation and membrane disruption.

When it comes to photodynamic therapy (PDT), the functionalization of gold nanoparticles can involve either directly attaching photosensitizers to the nanoparticle surface or encapsulating them within the nanoparticle matrix. This helps protect the photosensitizers from degradation and improves their pharmacokinetic properties. Moreover, targeting ligands can be added to ensure that the photosensitizer - loaded gold nanoparticles accumulate specifically in cancer cells, enhancing the specificity of PDT.

The combination of functionalized gold nanomaterials in photothermal and photodynamic cancer therapies shows great potential for developing more effective and less toxic cancer treatments. By precisely controlling the properties and targeting of the nanoparticles, these therapies can be customized for individual patients and different tumor types, offering new hope in the battle against cancer.

3) Functional gold nanomaterials have opened up new horizons in cancer biomedical diagnosis, providing sensitive, specific, and rapid methods for detecting cancer at an early stage and monitoring the progression of the disease. It can be used in electrochemical biosensors for the detection of cancer biomarkers in body fluids such as blood, saliva, and urine. The nanoparticles can be immobilized on an electrode surface and functionalized with antibodies or aptamers that specifically bind to the target

biomarkers. When the biomarkers bind to the functionalized nanoparticles, it causes a change in the electrical properties of the electrode, which can be measured and translated into a diagnostic signal. This approach offers the advantages of high sensitivity, low cost, and portability, making it suitable for point-of-care testing and early cancer screening in clinical settings.

4) The utilization of gold nanoparticles in genetic cancer therapy is a groundbreaking approach that holds significant promise for treating cancer at the molecular level. Gold nanoparticles can be designed to transport genetic substances like DNA or RNA into cancer cells. This enables the regulation of gene expression and the rectification of genetic mutations that are fundamental to cancer development. One of the tactics involves employing gold nanoparticles to deliver small interfering RNA (siRNA). siRNA molecules have the ability to specifically target and suppress the expression of oncogenes. These oncogenes are the genes that drive cancer cell growth and survival. By attaching siRNA to gold nanoparticles, they can be safeguarded from degradation in the extracellular environment and effectively transported into cancer cells. Once inside the cells, the siRNA is released and binds to the complementary mRNA of the oncogene, resulting in its degradation and the subsequent inhibition of protein synthesis. This method has shown great potential in preclinical research for treating various types of cancer, including lung cancer, breast cancer, and pancreatic cancer. Another application is the delivery of plasmid DNA that encodes tumor suppressor genes. Tumor suppressor genes play a vital role in controlling cell growth and preventing cancer. However, in many cases of cancer, these genes are mutated or rendered inactive. Gold nanoparticles can be utilized to introduce functional copies of tumor suppressor genes into cancer cells. This helps restore their normal function and trigger cell cycle arrest or apoptosis.

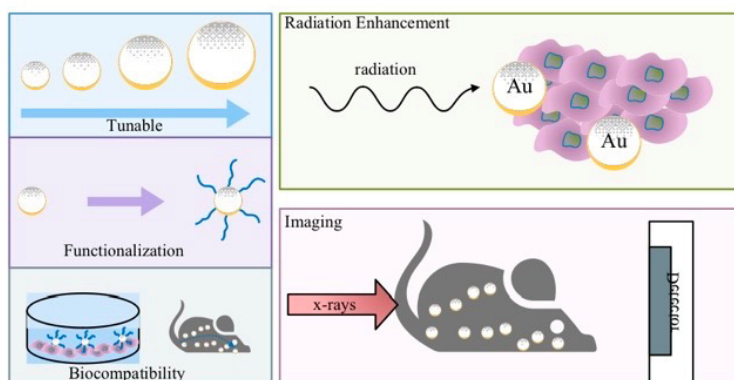


Figure 5. Radiosensitization with the aid of gold nanomaterials

5) Radiosensitization with the aid of gold nanomaterials is a burgeoning strategy in cancer treatment, striving to boost the efficacy of radiotherapy. Gold nanoparticles possess a distinct capacity to raise the local radiation dose inside cancer cells. This not only improves the therapeutic ratio but also has the potential to lower the radiation dose needed to control tumors.

The mechanism behind gold nanoparticles' radiosensitization is multifaceted. To begin with, owing to their high atomic number ($Z = 79$), gold nanoparticles can heighten the absorption of radiation energy. When exposed to X - rays or gamma rays, these nanoparticles produce secondary electrons via the photoelectric effect. These secondary electrons deposit extra energy near the nanoparticles. This increased energy deposition causes more severe DNA damage and greater cell death within tumor cells.

4. Risk of Gold Nanomaterial in Cancer Treatment

Gold nanomaterials have demonstrated substantial potential in cancer treatment. Nevertheless, their application entails several possible risks. A primary worry is the biocompatibility of gold nanomaterials and their toxicity to normal cells and tissues. Once introduced into the body, these nanomaterials might interact with healthy cells, disturbing normal physiological functions. Research

indicates that the size, shape, and surface coating of gold nanomaterials notably influence their toxicity. For instance, smaller nanoparticles may have a greater tendency to penetrate cells and inflict harm.

Another crucial factor is the degradation and elimination of gold nanomaterials post - treatment. After fulfilling their therapeutic function, it's vital to ensure their proper degradation and expulsion from the body. If gold nanomaterials build up in the body over time, they could lead to long - term health issues. The body's clearance systems, like the liver and kidneys, are pivotal in removing these nanoparticles. However, the efficiency of this process can differ based on the characteristics of gold nanomaterials. Moreover, gold ions that might be released during degradation require careful surveillance, as they too could have toxic effects. Comprehending and dealing with these potential risks is fundamental for the safe and effective use of gold nanomaterials in cancer therapy.

5. Conclusion

To sum it up, the investigation of gold nanomaterials in the field of cancer research and treatment has revealed numerous possibilities and progress. From the synthesis techniques that enable accurate control over their size and shape, to the various functionalization methods that boost their applications in targeted therapy, photothermal and photodynamic therapies, biomedical diagnosis, genetic therapy, and radiosensitization, gold nanomaterials have shown great potential. However, as we progress, it's crucial to deal with the existing challenges. More research is necessary to fully comprehend the long - term biocompatibility and potential toxicity of these materials in the human body, and to optimize their delivery systems for maximum effectiveness and minimum side effects. Besides, translating these promising laboratory results into clinical practice calls for collaborative efforts among scientists, clinicians, and regulatory agencies. With continuous innovation and a comprehensive understanding of the advantages and risks, gold nanomaterials have the potential to revolutionize the field of cancer treatment and diagnosis. They offer new hope for patients and lay the groundwork for more effective and personalized treatment approaches in the future.

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