

International Journal of Applied Pharmaceutics

ISSN-0975-7058

Vol 16. Issue 5, 2024

Review Article

EMERGING FRONTIERS: ADVANCEMENTS IN BIO-NONMATERIAL'S AND NON-INVASIVE STRATEGIES FOR COMBATING CANCER THROUGH PHOTO THERMAL THERAPY

YOKESH S.¹, TEEJESWARI R.¹, JALANIY V.¹, LOKESHVAR R.²

¹Saveetha College of Pharmacy, Saveetha Institute of Medical and Technical Sciences (SIMATS), Thandalam, Saveetha Nagar, Chennai-602105, India. ²Department of Pharmacology, Saveetha College of Pharmacy, Saveetha Institute of Medical and Technical Sciences (SIMATS), Thandalam, Saveetha Nagar, Chennai-602105, India ^{*}Corresponding author: Lokeshvar R.; *Email: lokeshvarr.scop@saveetha.com

Received: 11 Apr 2024, Revised and Accepted: 12 Jun 2024

ABSTRACT

Cancer remained a global health challenge in 2020 and claimed approximately 10 million lives, thus rendering it the main reason for demise. Photo Thermal Therapy (PTT) has emerged as a promising approach among the various cancer therapies. PTT offers several advantages over traditional treatments such as surgery, chemotherapy, and Radiotherapy due to its precise tumor targeting and reduced damage to healthy tissues. Photo Thermal Agents (PTAs) are central to PTT and selectively kill cancer cells by converting near-infrared light into heat. However, some PTAs exhibit toxicity and remain in the body's Reticulo Endothelial System (RES), limiting their clinical utility. To address this issue, scientists are looking at putrescible nano-photothermal compounds. This study talks about the effects of PTT and other cancer treatment techniques on patient health, stares at the process beneath PTT, and highlights recent bio-nonmaterial's utilized in PTT. It also looks at how low temperatures and brightly coloured light might be used to improve PTT efficacy.

Keywords: Bionanomaterials, Cancer, Photo thermal therapy, Photo thermal agents

© 2024 The Authors. Published by Innovare Academic Sciences Pvt Ltd. This is an open access article under the CC BY license (https://creativecommons.org/licenses/by/4.0/) DOI: https://dx.doi.org/10.22159/ijap.2024v16i5.51113 Journal homepage: https://innovareacademics.in/journals/index.php/ijap

INTRODUCTION

In this review, we explore the latest advancements in biononmaterial's and their application in Photo Thermal Therapy (PTT) for cancer treatment. The search criteria for the literature review included sources such as PubMed, Scopus, and Web of Science. Key words used were "PTT," "bio-nonmaterial's," "cancer treatment, "near-infrared light," "near-infrared-II materials," "nanoparticle toxicity," and "synergistic cancer therapies." The search was restricted to articles published from 2010 to 2024, ensuring a focus on recent developments. Peer-reviewed articles and studies involving preclinical and clinical trials were prioritized. Cancer treatment remains one of the foremost challenges confronting the current wellness sector [1]. With its alarming ephemerality rates and rapid spread, cancer poses a grave threat to individuals worldwide, claiming nearly 10 million lives in 2020 alone, equating to almost solitary ephemerality for every hexadsubject, with a projected worldwide fresh case count of 19.4 million [2]. Throughout the centuries, scientists have tirelessly pursued effective cancer therapies. While current treatments such as chemotherapy, radiotherapy, and surgery yield limited success and come with severe side effects, newer technologies have emerged, including gene therapy, immunotherapy, PTT, and Photo Dynamic Therapy (PDT) that have garnered popularity due to their promising attributes [3]. PTT has mainly garnered attention among these modalities for its heightened specificity, reduced unwanted reactions, potent anti – euplastic efficacy, and reverse-invasive nature [4]. PTT induces a photo thermal effect (light energy is absorbed and transformed into heat via this process), elevating the temperature of cancer cells and triggering organism death, as cancer cells are less heat-tolerant [3]. PTAs with small-scale engineering can actively and passively target tumors, resulting in improved PTT specificity. Only exposing cancer cells to the laser does less damage to healthy tissues. Curing cancer intends to eradicate tumors while preserving healthy cells [5, 6].

A diverse range of organic and nanoparticles made of inorganic material, including photo thermal agents, comprise nonmaterial's that include platinum, gold, and silver carbon-based nonmaterial's, near-infrared dyes, conductive polymers, soluble proteins and peptides, and melanin-like polymers. On the other hand, PTAs with near-infrared absorbance possess downsides such as suboptimal pharmacokinetics, low immunogenicity, elevated toxic effects, and

inflammatory side effects. They are also non-biodegradable. Investigating PTAs that exhibit strong clinical translational efficacy and optimize PTT therapeutic efficacy is imperative [7].

To solve the problem, scientists are creating biodegradable nonmaterials such as conducting polymers, melanin-like polymers, carbon-based tiny substances, and tiny molecular organic colors. These materials possess inherent biocompatibility and biodegradability, along with established degradation mechanisms and recognized metabolic pathways, making them promising candidates for enhancing the effectiveness of PTT. By leveraging the potential of these biodegradable nonmaterials, researchers aim to prevail over the constraints of recent PTAs and provide the most efficient, effective PTT treatments.

To ensure that PTAs used in clinical translation are successfully removed from the human body within a given timeframe, the Food and Drug Administration (FDA) of the United States emphasizes this matter. A clear advantage is provided by mechanisms that can stimulate the degradation of nonmaterial's [4]. Despite its considerable effectiveness in clinical therapeutics, PTT has drawbacks, especially when treating tumors at the periphery where light penetration is inadequate and partial removal of malignant cells results. This incomplete removal may facilitate tumor regret and metastasis to other body parts. Due to the previously mentioned limitations, PTT immunotherapy alone is currently frequently insufficient to achieve complete cancer eradication. Many complementary methods, including PDT, immunotherapy, gene therapy, and chemotherapy, are investigated to optimize PTT overall efficacy. These methods can lead to accurate tumor identification and management, providing a comprehensive plan to address a range of cancers and other conditions [8]. In addition to lowering the possibility of cancer cells spreading to all organs and unwanted reactions like inflammation, these combined approaches offer practical advantages over immunotherapy by reducing the chance of tissue injury, burns, and scarring from the heat near tumor locations [9]. Furthermore, nanozymes, a novel class of synthetic enzymes, have great potential.

Fan *et al.* made significant progress in creating a unique nanozyme with a characteristic yolk-shell structure in an innovative research study. With the help of this novel nanostructure, which combines a single gold nanoparticle core with porous hollow carbon shell

nanospheres, it is possible to see horseradish peroxides and oxidize-like functions in acidic environments. The gold nanoparticle core with porous hollow carbon shell nanospheres take full advantage of the photo thermal effect frequently used in tumor therapy by producing Reactive Oxygen Species (ROS) with remarkable efficiency, especially when exposed to 808 nanometer light. As demonstrated by extensive cellular and animal investigations, the combined effects of photo thermal therapy and light-enhanced ROS produce a highly effective tumor-eradication effect. These results highlight the potential for developing a novel tumor treatment strategy using nanozymes innate enzyme-like properties and photo thermal transformation. The research presents strong evidence for using nanozymes in catalytic PTT directed at tumors [10].

An ultra small platinum nanozyme and a carbon nanozyme generated from a Metal Organic Framework (MOF) were combined by Yang et al. to create a highly efficient nanozyme. By boosting catalane activity and converting hydrogen peroxide to oxygen, incorporating platinum nanozyme improved the effectiveness of PDT. Additionally, including the platinum nanozyme improved the carbon nanozyme's inherent photo thermal performance. The notable Inhibition of tumor growth observed in vivo with the platinum carbon nanozyme is attributed to the combined benefits of increased photo thermal characteristics and catalane-like activity [11].

Nie *et al.* To minimize bleeding after surgery, a device called Surgiflo based palladium-copper nanoclusters is intended for efficient insertion into different tumor cavities. This technology employs dispersed palladium-copper nanoclusters to improve the antitumor immune response and kill glioma cells by producing ROS following exposure to near-infrared laser irradiation [12].

Zhang *et al.* introduced metal active sites scattered over a substrate stabilized by coordinating with elements known as Single Atom Catalysts (SACs). This approach has produced exceptional catalytic

performance in various processes by maximizing the use of metal atoms. In disciplines like biology and nanotechnology, catalytic selectivity and activity are crucial factors to consider [13].

Under Zhang'sleadership, a research team has innovated a biodegradable hydro gel coupled with a bioadhesive to mitigate tumor recurrence post-surgery and mitigate skin damage from radiation therapy. This hydro gel facilitates oxygen release and drug delivery to combat cancer effectively, while the bioadhesive possesses antioxidant properties and fosters radiation wound healing. Animal trials have demonstrated the efficacy of this approach in suppressing tumor recurrence and mitigating radiation-induced skin heresy [14]. In this review, we have examined the most recent developments in biononmaterial used in PTT and the possible benefits of combining PTT with other therapeutic approaches. Additionally, we have provided insight into the role of mild hypothermia and Light Emitting Diode (LED) lights in enhancing the effectiveness of PTT.

Mechanism of photo thermal therapy (PTT) action

Over the past decade, there has been a remarkable surge in the utilization of PTT to eradicate solid tumors [15]. Towards PTAs at tumor locations, such as primary tumors and early local metastases, near-infrared light is directed. Heat is produced by this light-induced electron-to-electron relaxation within the PTAs, which causes hyperthermia and, ultimately, the death of cancer cells in the targeted areas (fig. 1) [16]. Several wavelengths are covered by PTAs, such as near-infrared-I (700-950 nm), near-infrared-II (1000-1350 nm), near-infrared-III (1600-1870 nm), and near-infrared-IV (2100-2300 nm). Because of their effectiveness as irradiation sources, PTT research has concentrated chiefly on lasers operating in the near-infrared-I window. Furthermore, because of its improved tissue penetration and advantageous safety profile, near-infrared-II, especially the near-infrared-II window (1300-1400 nm), is more effective than near-infrared-I [17]. PTT induces three primary mechanisms of cell death: necrosis, apoptosis, and necroptosis.

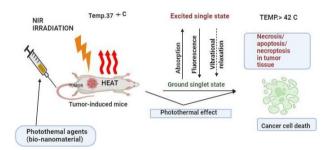


Fig. 1: Mechanism of photo thermal therapy (PTT) [70]

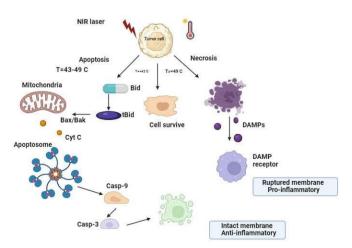


Fig. 2: Cell death mediated by photo thermal therapy (PTT) [70]

Necrosis is characterized by unplanned cellular demise, resulting in cytoplasmic enlargement, extensive organelle damage, and eventual

plasma membrane rupture [13]. This process exposes the extracellular environment to intracellular materials, including Damage Associated

Molecular Patterns (DAMPs), which can trigger detrimental immunogenicunfavourable mode of organism ephemerality [10]. Signals for phagocytosis are released during cell death, characterized by intact membrane integrity and membrane blebbing. Relative to the inflammatory reaction brought on by necrosis, it produces more significant results. It is possible to elicit necrosis without encountering resistance. Under ideal radiation circumstances, PTT can induce apoptosis. PTT induced apoptosis is facilitated mainly by the intrinsic route. It is interesting to investigate if the main pathway leading to cell death during apoptosis is the extrinsic or intrinsic pathway. Molecular signalling pathways linked to cellular responses to PTT offer insights into this matter. It is plausible that the primary mechanism of PTT induced cell death does not involve the extrinsic pathway, which is activated when specific ligands bind to "death receptors" on the cell surface. PTT probably causes membrane permeabilization and cytochrome c release from mitochondria by activating the intrinsic route. This outcome creates the apoptosome complex, which in turn causes a series of subsequent occurrences that culminate in cell death and phagocytosis by activating caspase-3, as depicted in fig. 2.

This is essential for highlighting the communication between the pathways leading to intrinsic and extrinsic apoptosis. When ligands engage death receptors, caspase-8 is activated and contributes to the cleavage of Bid into tBid. tBid then translocates to the mitochondria, releasing cytochrome c and activating Bax/Bak. This intricate interaction between pathways exemplifies the complexity of apoptotic signaling [15].

When cancer cells are subjected to high temperatures during PTT, typically exceeding 42 °C, they often undergo apoptosis or necrosis. The tumor cell ephemerality in PTT is known as necroptosis [18]. This mechanism shares similarities with a highly regulated cell death process and an uncontrolled, passive necrotic pathway. Reduced temperature can release heat-shock proteins, which prevent cancer cells from thermal burns. Irradiation with near-infrared light generates heat at 49 °C and below, which induces irreparable damage to tissue and necroptosis and cell death in cancer cells. Necrosis ensues when elevated temperatures disrupt the cancer cells membranes, releasing cytoplasmic contents and eliciting an inflammatory response. At 46 °C, protein denaturation and the disruption of cell membranes occur due to ischemia and micro vascular thrombosis, leading to cell death. Apoptosis, characterized by tightly controlled processes and the absence of inflammation, emerges as a suitable technique for eliminating cancer cells. PTT has acknowledged this distinction in cell death mechanisms for cancer treatment [14].

Bio-nanomaterials used for photo thermal therapy (PTT)

Excellent photostability, minimal harm to normal tissues, exceptional biocompatibility, and excellent photo thermal

conversion effectiveness at a particular wavelength is the ideal PTAs characteristics. Fig. 3 depicts various types of PTAs commonly employed, including organic nonmaterial's like conductive polymers and near-infrared dyes, noble metal nonmaterial's such as gold nanorods and gold nanospheres, and metal compounds like iron, copper, and molybdenum [19]. In recent years, a range of inorganic excellentphotostability, minimal harm to normal tissues, exceptional biocompatibility, and excellent photo thermal conversion effectiveness at a particular wavelength are characteristics of the ideal PTAs. PTAs has been synthesized and investigated in both in vitro and in vivo studies due to their surface plasmon resonance properties, which confer excellent near-infrared light absorbance [20]. However, despite their promising therapeutic efficacies, the drawback of these nonmaterials lies in their toxicity and nonbiodegradability, limiting their clinical translation and therapeutic applications [21].

In comparison, organic nonmaterials demonstrate higher biocompatibility [22]. To address this issue, various biononmaterials have been synthesized to enhance the photo thermal effect while ensuring biocompatibility, biodegradability, low cytotoxicity, and rapid clearance from the body. This biononmaterial's are discussed in the following section and are summarized in table 1 [70].

Carbon-based nanomaterials

Carbon nanotubes (CNTs)

These are cylindrical sp2 hybridized monolayer sheets of carbon atoms [23]. Due to these unique attributes, Carbon Nanotubes (CNTs) are currently at the forefront of scientific investigation. Serving as plasmonic nanoparticles, CNTs can infiltrate cells due to their remarkable thermal conductivity [42]. The thermal conductivity of CNTs offers several advantages, including even distribution of generated heat throughout tumor cells and confinement of heat exclusively to the targeted area, thereby reducing toxicity to off-target tissues. Furthermore, Magnetic Resonance Imaging (MRI) temperature mapping can verify targeted tumor tissue volume [23].

Nanotubes come in two primary varieties: single-walled and multi-walled. As the name suggests, Single-Walled Carbon Nanotubes (SWCNTs) consist of one layer forming a cylinder while the multi-walled nanotubes comprise several cylindrical layers each having a diameter greater than the other. The chemical bonding in the tubes is best described by orbital hybridization (sp2-hybrid carbon atoms) which accounts for the unique strength of CNTs [102].

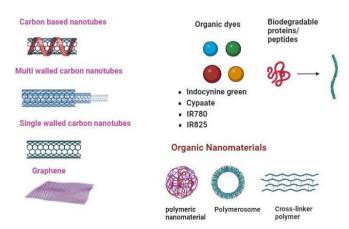


Fig. 3: Different biomaterials used for photo thermal therapy (PTT) [70]

Single-walled carbon nanotubes (SWCNTs)

Graphitic helical molecules, known as Single-Walled Carbon Nanotubes (SWCNTs), possess remarkably robust mechanical and physical properties [24]. Studies conducted by Liu *et al.* concluded that SWCNTs exhibit excellent biological stability, low toxicity, and

favourable water solubility [43]. Liang *et al.* similarly found that SWCNTs demonstrated high strength and minimal cytotoxicity in both *in vitro* and *in vivo* experiments. Incorporating Poly Ethylene Glycol (PEG) into SWCNTs prolongs their circulation time in the bloodstream. In a study by Li *et al.*, the addition of cyanine 5.5, which absorbs light at 808 nm, enabled cyanine 5.5 coupled SWCNTs to

mediate PTT in mice, resulting in systemic tumor ablation and improved therapeutic outcomes [99]. Virani et al. employed bladder tumor-specific SWCNTs followed by near-infrared light to treat superficial cancers and stop tumor recurrence. The bound nanotubes that are led to the tumor site by phosphatidylserine and annexin V are the main targets of the heat-based treatment [45]. Li et al. highlighted the SWCNTs exhibit remarkable Photothermal Conversion Efficiency (PCE), excellent water solubility, bio stability, and minimal toxicity. They can be coupled with dye and antibodies to give dye imaging-guided cytotoxic PTT that targets pancreatic cancers [46]. Mei et al. synthesized selenium based CNTs, which, when exposed to radiation, produce ROS and cause tumor cells to undergo apoptosis. By modifying signal transduction pathways, the nanosystem prevented Triple Negative Breast Cancer (TNBC) cells from migrating and invading [23].

Mckernan et al. introduced a novel therapeutic strategy for metastatic Epithelial Mammary Tumor 6 (EMT6) breast cancer in

BALB/c jackson mice that are syngeneic,integrating PTT with targeted SWCNTs and immunostimulation using a checkpoint inhibitor. This newly formulated approach enhanced the abscopal effect, relying on the cytotoxic T-lymphocyte associated antigen 4 blockade, showcasing remarkable synergy. As a result, there was a significant increase in the living probability of mice, reaching 60%. Moreover, this treatment regimen boosted the population of CD4+helper and CD8+cytotoxic T-cells, thereby enhancing the activity of anti-tumoral effector cells and inhibiting tumor spread [47]. Chen et al. developed an amylose derivative conjugated to modified SWCNTs. The amylose derivatives containing poly (L-lysine) dendrons complex exhibits excellent water dispersion stability and demonstrate high p - Deoxyribo Nucleic Acid (DNA) transfection capacity. In both in vivo and in vitro studies, Tumor Necrosis Factor (TNF)/amylose derivatives containing poly (L-lysine) based SWCNTs effectively inhibits tumor growth and prevents metastasis. The potent antitumor effects of near-infrared irradiation suggest its potential utility as a cancer therapy [24].

Table 1: An assortment of biological-tiny materials used in photo thermal therapy (PTT)

S. No.	PTAs	Sample	Carcinoma types	Curative trace	References
1.	Mxene microcapsules	Titanium carbide with Mxenes	Ductal carcinoma	The growth of cancer cells is reduced	[4]
2.	Amino acid-based Nanomaterials	Paclitaxel-ICG-Human Albumin Serum (HAS) nanoparticles	4T1 Epidermal cysts	Neither carcinogenicity Nor tumor development	[37]
3.	Nanomaterials based on carbon				
a.	CDs	Lecithin-coated red fluorescent CDs	Ductal carcinoma	Enhanced rate and extent of drug absorbed in the systemic circulation and Biological action	[26]
b.	CNTs				
i.	MWCNTs	MWCNTs with PEG CNTs onto PEG	Bronchogenic Cancer Cardiac sarcomas	High photo thermal treatment effectiveness and firm tumor targeting ability After PTT, there Is reduced Cytotoxicity and a lesser-Sized tumor.	[25] [21]
ii.	SWCNTs	Selenium based CNTs TNF/amylose derivatives containing poly (L-lysine) based SWCNTs	TNBC Any kind	The formation of ROS, Leading tumor Cells to Undergo death A high level of Stability of Water Dispersion and p-DNA transfected capacity	[23] [24]
c.	Graphene				
i.	GO	GO nano complexes based doxorubicin-folic acid nanomaterial's	Ductal carcinoma	Excellent PCE	[29]
ii.	GQDs	9T-GQDs	Ductal carcinoma	More properties of both light and heat	[32]
iii.	GN	Polydopamine doped mesoporous silica-coated reduced rGO	Hepatoma	More medication will be let out at the site of cancer cells.	[27]
iv.	rGO	Plant extract-based rGO GO/BaHoF5/PEG Tamoxifen nano rGO	Esophageal adenocarcinoma cell lines Adenosquamous Carcinoma Ductal carcinoma	Cancer cells Destroying May utilize EPR impact to get into spots Effective tumor killing action	[31] [28] [30]
4.	Small Molecular Organic Dyes (SMOD)				
a.	Infrared-780	TM 815	Hepatocellular carcinoma	To image deep tissues	[2]
b.	ICG	Monomethoxy PEG-luteolin- bortezomib@ ICG	Bowel carcinoma	Destroying of cancer cells	[33]
c.	Cypate	Silicon dioxide-ammonium based ICG nanoparticles	Hepatocellular carcinoma	Considerable tumor cell death	[34]
5.	Nuclear Magnetic Resonance Spectroscopy (NMRs) based on polymers				
a.	Polymer(melanin)	PEG ylated nanoliposomes containing melanin	Basal cell carcinoma	Better Bioabsorbable	[41]
b.	Using conducting polymers				
i.	Pyrrole polymer	Polypyrrole-poly (ethylene imine)- silk nanoparticles	Papillary carcinoma	Suppression of lymph cancer cell spread to other organs	[40]
ii.	Polyaniline	Gold nanostar@polyaniline	4T1 cells	Magnificent PCE	[39]

Multi-walled carbon nanotubes (MWCNTs)

Multi-Walled Carbon Nanotubes (MWCNTs) exhibit diameters ranging from tiny to very small nanometers. They are increasingly recognized as promising candidates for applications in biological imaging, photo thermal tumor ablation, and tumor drug delivery [50]. Compared to other materials like SWCNTs, MWCNTs are anticipated to absorb a significantly more significant amount of near-infrared radiation, potentially leading to tumor thermal

ablation through localized heat generation upon near-infrared exposure. Cys-Arg-Glu-Lys-Ala peptide, PEG, MWCNTs, fibrin, and near-infrared irradiation are the therapeutic payloads of Zhang *et al.*'s drug delivery system, MWCNTs with PEG. It demonstrated efficacy in photo thermal therapy and had good tumor targeting [25]. Sobhani *et al.* covered MWCNTs with PEG and oxidized them to improve their water permeability. Tumor development and cytotoxicity decreased in CNTs onto PEG after PTT. A high-temperature conductivity and absorbance in the near-infrared range

have been achieved by combining CNTs with gold nanoparticles. The authors found MWCNTs with gold nanoparticles effective in cancer treatment [42]. These findings underscore the promising clinical and therapeutic potential of nano-agents despite the ongoing uncertainty regarding their long-term safety [21].

Carbon dots (CDs)

Zero-dimensional Carbon Dots (CDs) are nonmaterial's with exceptional optical characteristics, biocompatibility, and little environmental effect. Because they can convert light into heat, they have significant promise for various applications, including photo thermal applications [48]. Shinde et al. demonstrated the utilization of Clitoria ternatea leaves in synthesizing hydrophilic lecithin-coated red fluorescent CDs with enhanced bioavailability and theranostic capabilities. With a hydrodynamic diameter of roughly 210 nm, soy lecithin-coated red fluorescent CDs are nanoparticles suitable for bioimaging and cancer therapy. They have assured PTT agents, which can cause significant cell death in 4T1 cells and damage neoblood vessels in Chorio Allantoic Membrane (CAM) assays [26].

Graphene

Recent interest in graphene-based nonmaterials has been driven by their unique structural, optical, thermal, electrical, and mechanical properties [49]. Graphene, a two-dimensional nanosystem of sp2-hybridized carbon atoms arranged in a hexagonal pattern, has garnered significant attention [43]. Because of its biocompatibility, Graphene Oxide (GO) and reduced Graphene Oxide (rGO) are widely used in PTT [23]. Because of its intense near-infrared light absorption, graphene is a suitable nanoagent for PTT [21].

Graphene nanosheet (GN)

The photo thermal properties of GO nanosheets exhibit remarkable performance and are characterized by a broad absorption spectrum. Additionally, the ample surface area of GO renders it a promising platform for drug loading. Dopamine hydrochloride was utilized by Liu *et al.* to generate polydopamine doped mesoporous silica-coated rGO. The photo thermal conversion efficiency and the release of medication at the tumor location are improved by this substance [27].

Furthermore, Wang Yijuan *et al.* conducted docking study utilizing methods of Monte Carlo, revealing an uneven distribution of nanosheets on the surfaces of cancer cells. This uneven distribution led to a rapid temperature increase, facilitating tumor cell death [100]. In another study by Li *et al.*, manganese dioxide nanoparticles were incorporated into the rGO sheets mixture. The solid physical bond between methylene blue and doxorubicin with GO increased drug release at high temperatures [51].

Graphene oxide (GO)

Graphene Oxide (GO) possesses several oxygen-based functional groups. These significantly reduce graphene's hydrophobicity [43]. Many GO small particle varieties utilized in photo thermal therapy are outlined in table 2 [70]. Leveraging extensive surface area for drug binding through hydrophobic, H-bonding, π - π , and electrostatic interactions [23]. By loading doxorubicin onto the nanocomposite, Liang et al. created GO nano complexes based doxorubicin-folic acid nanomaterial's. As a result of its large surface area, target selectivity. drug loading capacity, PCE, and photostability when exposed to radiation at 808 nm, this formulation works well [28]. Chang et al. engineered a flexible nano platform by binding barium hole-doped iron oxide fluoride 5 nanoparticles to GO. Leveraging the Enhanced Permeability and Retention (EPR) effect, the biocompatible GO/barium hole-doped iron oxide fluoride 5/PEG nanocomposite infiltrates lesions efficiently. This nano platform can be loaded with Heat Shock Protein 90 (HSP 90) NVP-AUY922 inhibitor to combat cancer and perform PTT upon irradiation with an 808 nm nearinfrared laser [29]. Qi et al. developed a Schiff base-cross-linked hydro gel using Carbonyl Methyl Chitosan (CMC) and aldehydemodified PEG. This injectable and self-healing hydro gel was designed to carry GO and nano hydroxyapatite inhibitors in needles, enhancing tumor diagnosis accuracy while chemotherapeutic side effects. The hydro gel exhibited self-healing properties, improved injectability, and a porous structure [52]. Baipaywad et al. engineered a controlled in vivo drug delivery system by combining GO and poly N-isopropyl acrylamide in a nano hydro gel formulation. By strengthening PTT effectiveness in nano gel systems, GO and poly N-isopropyl acrylamide generated doxorubicin when exposed to near-infrared light [53].

Table 2: The compilation of photo thermal therapy (PTT)-utilized graphene oxide (GO) nanomaterial's

S. No.	Formulation	Curative marks	Reference
1.	GO and poly N-isopropyl acrylamide	Less toxic against tumors and biologically compatible	[53]
2.	PEG-CMC/hydroxyapatite/GO	Completely stop the cancer cell proliferation	[52]
3.	GO/BaHoF5/PEG	Positive PTT impact on cancer	[29]
4.	GO nano complexes based doxorubicin-folic acid	Outstanding possibilities for cancer treatment applications	[28]
	nanomatorial's		

Reduced graphene oxide (rGO)

Due to its high PCE in the near-infrared range, enormous surface area, and groups of molecules suitable for small interfering Ribo Nucleic Acid (RNA) and ligand attachment, reduced Graphene Oxide (rGO) is a promising 2D carbon nanomaterial for photo thermal therapy [54]. Regarding PCE, rGO demonstrates a two-fold improvement over GO as a PTT agent [29]. Various types of rGO utilized in PTT are outlined in table 3 [70]. Zhang et al. devised a straightforward method for putting medications into rGO that is nanoscale for breast cancer PTT, employing tamoxifen as the medication, resulting in significant tumoricidal activity with the combined tamoxifen nano rGO composition [30]. Gai et al. demonstrated that rGO derived from synthetic plant extracts exhibits minimal cytotoxicity towards cells and can be directly employed for PTT without additional surface modifications, effectively destroying Esophageal adenocarcinoma cell lines in vitro [31]. A tiny carrier combination for synergistic chemo PTT was created by Liu et al. (rGO@mesoporous silica), featuring an outer layer composed of mesoporous silica and an inner layer of rGO. This structure facilitates the easy release of doxorubicin into acidic environments, serving as both a near-infrared PTT agent and a pHinduced drug nanocarrier [55]. Lima-Sousa et al., in collaboration with other researchers, developed poly dopamine rGO by utilizing rGO as a reducing agent and dopamine. According to a study, poly dopamine rGO indicated considerable near-infrared spectrum absorption and kept distributed all over the cells. Thiol-terminated poly(2-ethyl-2-oxazoline) was attached to enhance its stability and characteristics [42]. While remaining stable and compatible with organisms, this boosted poly dopamine rGO version exhibited intriguing properties. U87 glioblastoma cells showed uptake of this nanosystem. They developed an infrared-780/rGO/Hyaluronic Acid (HA) pH-responsive nano system loaded with doxorubicin for multimodal cancer treatment [chemotherapy/PTT/PDT] [56].

Graphene quantum dots (GQDs)

Solid quantum dots, abbreviated as Graphene Quantum Dots (GQDs), are nanostructures known for their photoluminescence and prominent edge effects. Leveraging an near-infrared laser, GQDs have found extensive application in PTT [49]. Because of their low level of toxicity and outstanding absorption in the near-infrared-II region, GQDs have tremendous potential for application in photo thermal therapy. Robust magnetic field tuning of hydrogen peroxide breakdown was achieved by Liu et al. [32].

Table 3: The inventory of photo thermal therapy (PTT)-used reduced graphene oxide (rGO) nanoparticles

S. No.	Formulation	Curative marks	Reference
1.	Infrared-780/reduced rGO/HA	For enhancing combined activity for [chemotherapy/PTT/PDT] treatment	[56]
2.	Poly dopamine rGO	PTT For the utilization in Ductal cancer	[57]
3.	rGO @mesoporous silica	For cancer treatment, it was the suitable platform	[55]
4.	Reduced nano GO	Kill Oesophageal adenocarcinoma cell lines	[31]
5.	Nano-rGO	Destroying of cancer cells	[30]

Small molecular organic dyes

Molecular dyes exhibit exceptional characteristics concerning surface modification and compatibility with biological systems. Tailoring molecular dyes to meet specific absorption wavelength requirements, particularly for targeted loading into tumor cells, can be achieved through seamless reshaping [58].

Indocyanine green (ICG)

Indocyanine Green (ICG) has gained approval from the USFDA as a clinical diagnostic photosensitizer. As ICG comes into interaction with near-infrared laser radiation, it may trigger hyperthermia and generate ROS. However, due to its rapid clearance rates and poor stability, utilization in PDT is constrained. For a significant load of ICG, nanocarriers that include polymeric nanoparticles, calcium phosphate nanoparticles, and mesoporous silicon nanoparticles are being investigated [43]. Qing et al. produced monomethoxy PEGluteolin-bortezomib@ ICG, a pH-responsive gel incorporating chemotherapy medications and PTT/PDT to manage colorectal cancer. Through the synergistic action of bortezomib and ICG, this method successfully eliminated the tumor cells in both in vitro and in vivo trials [33]. Prussian blue nanoshells facilitated the absorption of ICG particles, forming fine aggregates that enhanced the cellular of drugs. This combination exhibited improved histocompatibility, hemocompatibility, increased apoptosis, reduced toxicity, and effective tumor destruction [59]. ICG is a robust photosensitizer that booststhe production of ROS.

Additionally, its unique ability to absorb near-infrared light enhances the photo thermal effect, contributing to the degradation of tumor cells [60]. Wang and colleagues altered the surface potential of silicon dioxide-ammonium nanoparticles by incorporating ICG. This modification enhanced the targeting capacity of silicon dioxide-ammonium based ICG nanoparticles, making them stable enough to execute PCE. These silicon dioxide-ammonium based ICG nanoparticles absorb more light in the near-infrared spectrum after coating the ICG surface. According to this study, these nanoparticles significantly kill tumor cells [34]. To better the visible properties of ICG and enhance its uptake in 4T1 memory malignancies, Miranda *et al.* merged ICG to liposomes containing 1,2-dioleoyl-3-trimethyl ammonium propane. They examined the continued effectiveness of the combined formulation by using fluorescence and photoacoustic imaging approaches [61].

Cypate

Cyclapate, an indocyanine green derivative containing bis-carboxyl groups, exhibits heat production and fluorescence upon exposure to near-infrared radiation. Yin and colleagues developed Cypate-Poly Methyl Methacrylate (PMAA)-iron based Mesenchymal Stem Cells (MSCs) by coating PMMA nanoparticles with membranes derived from MSCs loaded with Fe and Cypate. This novel formulation demonstrates high PCE *in vitro* and *in vivo*, along with excellent tumor accumulation and stability, making it suitable for diagnosing lung cancer. Moreover, Cypate-PMAA-iron based MSCs serve as a platform for MRI bimodal imaging, fluorescence, and PTT enhanced radiotherapy [35].

Infrared-780

Concerning ICG, infrared-780 dye is brighter but mixes more slowly in water and exits the body more quickly. The stability of infrared-780 dye in water may be boosted by wrapping it in micelle nanoparticles [48]. Biomolecule extracts' lipophilicity restricts the possibility of enhancing the therapeutic potential of several beneficial plants. In a study, Pemmaraju *et al.* used poly D-lactic co-glycolic acid to

encapsulate and distribute a hydrophobic extract from Anthocephalus cadamba. They added infrared-780 dye to the mixture as well. To create a cancer treatment that works, this strategy sought to increase the cells' absorption and the bioavailability of the Anthocephalus cadamba extract. HD of 177.3 \pm 85.5 was observed by infrared-780/Anthocephalus cadamba polymeric nanoparticles. At 20µg/ml concentrations, polymeric nanoparticles demonstrated enhanced cytotoxicity and selectivity for tumor cells, resulting in 53.2% of B16 cell lines collapsing [36]. Tutty, in addition. No cytotoxicity or damage to DNA was found when using liposomes loaded with the dye infrared-780 (LipImage (TM) 815) for deep tissue imaging [2].

Numerous other dyes besides infrared-780 also produce photo thermal effects, such as Pebam. In an investigation of malignant pulmonary cells, tiny polylactic acid particles comprising infrared-775 and an ethyl acetate extract of Terminalia chebula, a plant, were used. The results showed that near-infrared radiation elevated cell death decreased HSP70 and poly adenosine diphosphate ribose polymerase, and upregulated Phosphorylated Histone H2A Variant X [62]. A chitosan-coated Niosome has been proposed by Jogdand *et al.* as an approach to giving infrared-806 dye to inhibit the synthesis of mucin in breast cancer patients. When near-infrared radiation was utilized on Michigan Cancer Foundation-7 (MCF-7) cells, the nanosystem was activated and ablated cells by up to 93% [63].

Proteins/Peptides-based nanomaterials

While they are dissolved in water and exhibit an elevated level of biocompatibility, peptides and amino acids are excellent carriers for medicinal or diagnostics elements [7]. They may act as biotemplates for the synthesis of nanoparticles. Biomolecular hydrogels can target the delivery of medications, reduce cytotoxicity, and enhance drug release [64]. Subcutaneous carcinomas can now be eliminated with stable nanoparticles that do not trigger toxicity or stimulate tumor growth [37].

Polymer-based nanomaterials

A polymer of tiny particles was developed by Mudigunda *et al.* to treat retinoblastoma. Palladium(II) phthalocyanine nanoparticles were exhibited by Appidi *et al.* to induce DNA damage and cell division in tumor cells[66].

Conducting polymers

Conducting polymers, such as polyaniline and polypyrrole, have found widespread use in biomedical and bioelectronic applications. Due to their favourable optical absorption quality, they are particularly well-suited for PTT. Although conducting polymers are considered more biodegradable and less costly, they are often used as elimination agents in photo thermal treatments [67]. Poly-(3,4ethylene dioxy thiophene):poly-(4-styrene sulfonate), a conducting polymer made from polythiophene, has been paired with PEG to improve biocompatibility. It can also be used as a polymer PTAs for cancer PTT [68]. Mice with 4T1 tumors were also employed to test double-acceptor polymers with conjugation [Tryptophan Tryptophyl Quinone (TTQ) and Diketo Pyrrole (DPP)], enhancing photo thermal capacity and generating resilient near-infrared-II F1 signals. Therefore, they have vast potential for tumor eradication[69]. The work of Liu et al. and Chen et al. exemplify the possibility of conducting polymers in cancer therapy. Poly-(3,4-ethylene dioxy thiophene):poly-(4-styrene sulfonate), PEG and polymer-1 nanoparticles, in particular, have demonstrated excellent biocompatibility, photostability, and photo thermal capacity, which make them attractive candidates for future research and development in this field.

Polyaniline

The first documented photo thermal anticancer agent based on polymers is called polyaniline. A promising hydro gel Guar Gum (GG) based iron with polyaniline was created by Huang et al. Borax) through the doping of iron with polyaniline linked to GG moieties through borate/diol linkages. Guar gum chains were first used in this investigation as the main building component. GG based iron with polyaniline-borax demonstrated amendable sol-gel transformation, good injectability, and quick self-healing characteristics in response to heat or pH stimulation. Within the hydrogen Peroxide enriched tumor microenvironment, iron-doped polyaniline produced free OH radicals. The astonishing photo thermal characteristics and regulated release of pharmaceuticals in the near-infrared triggered range exhibited by hydrogels enable the establishment of combination PTT, radiation therapy, or PDT for in vitro and in vivo melanoma contexts [38]. An efficient polyaniline core-shell was created by Wang et al. Nanomaterial with an exceptional PCE administered at low doses for Photo Acoustic Image (PAI)-guided photo thermal therapy. HA modifies this PTAs without compromising its photo thermal effectiveness. Gold nanostar probe hydrogels demonstrate outstanding biocompatibility in PAI-guided PTT to removetumours [39].

Polypyrrole

Organic polypyrrole properties include improved biocompatibility and a high PCE. The polypyrrole-cantharidine was created by Cheng et al. Matériaux Institut Lavoisier-100 (MIL-100) based plasmonic core multimodel nanoparticles with coated macrophage cell membranes) successfully raised the PTT-based Fenton reaction rate. As a result, iron ions and polypyrrole cantharidine are released into the plasmonic core multimodel nanoparticles [101]. Near-infrared photosensitivity is present in polypyrrole. Polypyrrole-poly (ethylene imine)-silk nanoparticles were created by Xu et al. The combination of photo thermal and gene therapy was evaluated with this formulation. It reduced the invasive ablation of the malignancies and inhibited lymphatic metastasis [40].

Melanin-like polymers

Antioxidant, radio-resistant, and anti-euplastic, this polymer is present in many plants, animals, and human tissues. Furthermore, melanin has intense near-infrared light benefits PTT [68]. In a biocompatible approach, Zhang *et al.* revealed that melanin PEG ylatednanoliposomes had good biocompatibility and clinical translation [41]. The prospective use of polyphenol-based PTAs and melanin-like poly dopamine substances as therapies for cancer is

being investigated due to their sustainable production and biodegradable and biological compatibility [9].

Microspheres of MXene for photo thermal therapy (PTT)

A novel two dimensional (2D) transition metal called MXene [1] was discovered in 2011 by Naguib *et al.* The parent metal, aluminium, carbon or nitrogen phase is printed to remove the element of the A-group, aluminium. As a result of their outstanding photo thermal conversion efficiency, enormous surface area, and simple functionalization, MXenes are intriguing substances for cancer research [3]. In breast cancer models, the full potential for tumor cells to respond to PTT has been explored; this has resulted in a reduction in the lifespan of cells and increased reactive oxygen species. In bioprinted models, PTT-mediated MXenes also impeded the growth of cancers [4].

Naphthalocyanines (Nanonaps) and phathalocyanine

Zhang *et al.* created nano-formulated naphthalocyanines, or nano-naps, exhibiting remarkable contrast in the near-infrared range for cancer theranostics. Multipurpose uses for nanonaps include lymphatic mapping and whole-tumor PAI. Additionally, they have inherent qualities that make them appropriate for Positron Emission Tomography (PET). A topical up conversion skin cream was developed using nanonaps for PTT against tumors. Molecules such as phthalocyanine and porphyrin have potential uses in imaging and medicine [71].

Phthalocyanine is an organic PTAs that is highly flexible and environmentally acceptable. Its chemical structure is well-defined and exhibits significant absorption in the near-infrared spectrum [72]. It made a novel cruciform phthalocyanine pentad that forms water-soluble nanoparticles with unique absorption capabilities in the near-infrared region. With a phenomenal PCE of 58.3%, these nanoparticles produce a robust photoacoustic signal [73].

Combinatorial effect of photo thermal therapy (ptt) with cancer therapy

PTT has become increasingly popular in recent decades due to its increased therapeutic efficacy. PTT still has a lot of drawbacks, though. Factors that prevent its use in many clinical situations (such as high temperature, a shallow depth of light penetration, thermal injury to normal tissues, and heat-shock response) (fig. 4) [74]. To circumvent these constraints to deliver effective cures for cancer, combination therapy-which combines concurrently co-targeting substances or combining medical treatments in various forms-is needed. It is true that combination therapy significantly improves the prognosis for a wide range of illnesses [33].

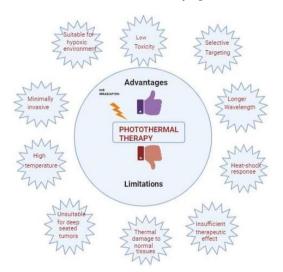


Fig. 4: Advantages and limitations of photo thermal therapy (PTT) [70]

Photo thermal therapy (PTT) combined with chemotherapy

For advanced-stage tumors, chemotherapy is an effective cancer treatment. For many cancer patients, it extends their life expectancy. This therapy is difficult to use, nevertheless, due to issues like limited drug penetration in tumor locations, significant cytotoxicity, and non-specific distribution. Better outcomes have been achieved by adapting a combination of chemotherapy and PTT to get around these constraints [16].

Synergistic PTT/chemotherapy action were created by Liu *et al.* Epoxide Iron(III) Chloride Nanoparticles (EICN) is a tremendous therapeutic drug for chemotherapy/PTT because of its dual-acid/near-infrared-responsive qualities. EICN is very stable and has a diameter of 90 nm [75]. Li *et al.* developed a novel nano agent that delivers the chemotherapeutic medication Paclitaxel using polymeric nano micelle-based technology. When this nano agent is exposed to an 808 nm laser, site-specific hyperthermia is quickly achieved. It is an efficient tumor treatment because of the temperature increase and site-specific medication delivery [76].

A novel approach utilizing chemo-PTT generated by near-infrared light is used to manufacture doxorubicin-Bovine Serum Albumin (BSA)– rGO nanosheets. They have shown promise in addressing brain tumor cells. The ensuing nanosheets were evaluated with X-

ray Photoelectron Spectroscopy (XPS) and an Ultra-Violet (UV)-visible spectrophotometer [77].

Photo thermal therapy (PTT) combined with immunotherapy

While PTT is less intrusive on surrounding tissue, its effectiveness in stopping tumor recurrence is short-lived and results in cytotoxicity. PTT agent biodegradation is sluggish [78]. Immunotherapy is a potentially effective treatment method that stimulates the host's immune system to fight the tumor. Restricted immune cell penetration, restricted immunity, alteration rate, and a wrong host immune response are some of the disadvantages of immunotherapy. For it to fight cancer, a combination regimen, including PTT-induced immunotherapy, releases immune responses that are adaptive as well as innate (fig. 5) [16].

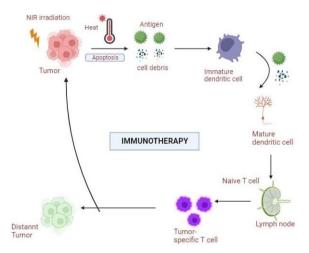


Fig. 5: Photo thermal therapy (PTT) combined with immunotherapy [70]

Immune checkpoint inhibition paired with photo thermal therapy leads to focused cytotoxicity and improved immunogenicity, improving cancer patients' clinical outcomes [79, 80]. In addition, gellan gum hydro gel, polyoxometalate, and localized ablative immunotherapy have shown positive results in boosting the anticancer defenses against breast tumors [78].

Photo thermal therapy (PTT) combined with radiotherapy

For improved clinical translation outcomes, Radiotherapy is a frequently employed approach combined with radio sensitizers [81]. Some restrictions exist, like weakening the ability to destroy and arming the surrounding solid tissues. Consequently, to maximize radiotherapy effectiveness, it is imperative to combine it with other medications, including PTT (fig. 6) [82].

Using Poly (thiodi-ethylene malonate) and PEG), both packed with radiosensitive Suberoyl Anilide Hydroxamic Acid (SAHA), Liu *et al.* constructed a nano-phototherapeutic system. The equipment demonstrated its capacity to carry out morphological changes and generate therapeutic effects in responses to near-infraredirradiation and X-ray radiotherapy. As compared with radiation treatment alone, a combination of therapy diminished the ability to survive breast cancer cells [81]. Radio therapy, PDT/PTT effects, near-infrared imaging, and preferential tumor accumulation are all displayed by the multifunctional bioactive small molecule infrared-83. It was built with intrinsic tumor penetration and phototherapeutic efficacy using 2-nitroimidazole, a radio sensitizer, in the core of heptemethine cyanine dyes [82].

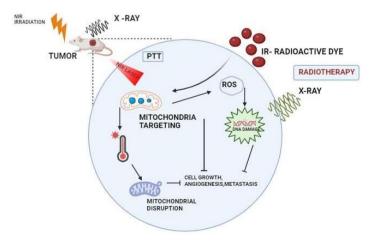


Fig. 6: Photo thermal therapy (PTT) combined with radiotherapy [70]

Photo thermal therapy (PTT) combined with gene therapy

Gene vectors improve gene therapy therapeutic application of nucleic acids DNA, small interfering RNA and micro RNA [83]. The negative aspects of gene Therapy include allergic reactions, poor clinical trials, and safety issues. Odda et al. utilized an unusual photo thermal nanocarrier to combine PTT in gene therapy. Small interfering RNA-B cell lymphoma 2 was employed to capture and load the surface-modified iron oxide based poly (3,4-ethylene dioxythiophene) nanoparticles into breast carcinoma cells. Compared to other iron oxide based poly (3,4ethylene dioxythiophene) nanoparticles, this core-shell surface-modified iron oxide based poly (3,4-ethylene dioxythiophene) nanoparticles have outstanding demonstrated intense near-infrared irradiation. biocompatibility, and water dispersibility. They demonstrated improved photostability and PCE (n=54.3%) in the area. The combination treatment of gene therapy and PTT, as opposed to either therapy alone, showed superior tumor cell death, according to the results [84]. Liu and colleagues generated HA-coupled. It Is anticipated that this will accelerate the body's nanoformulation circulation time. These composites showed improved targeting of tumor tissues, stability, biocompatibility, and a longer retention time in the Physique [85].

Photo thermal therapy (PTT) combined with photo dynamic therapy (PDT)

In recent years, PDT has become a very appealing alternative therapy. When exposed to laser light, it produces ROS by employing photosensitizers to impact the cancer cells' death [44]. Numerous benefits of this therapy include its excellent selectivity, reduced invasiveness, exceptional healing abilities, and reduced adverse effects. However, tumor hypoxia or reduced tissue penetration limits its application in clinical translation. The combinatorial impact of PTT with PDT yields the intended result and fixes the problem [16]. Fig. 7 explains the PTT and PDT combinatorial mechanism.

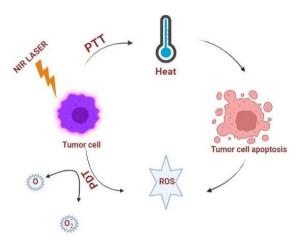


Fig. 7: Photo thermal therapy (PTT) combined with photo dynamic therapy (PDT) [70]

Spinacia oleracea co-assembled on a polydopamine core was the source of the multifunctional chlorophyll-rich fluorosomes that Rajalakshmi *et al.* proposed to enhance the process of photo thermal transmission. Using a combination of PDT and PTT, Spinacia oleracea co-assembled on a polydopamine core nanoparticles were assessed the proliferation of cancer cells under normoxic and hypoxic conditions. Synergistic PTT and PDT improved physiological stability and tumor regression in Spinacia oleracea co-assembled on a polydopamine core nanoparticles, which measured 176±10 nm. When Spinacia oleracea co-assembled on a polydopamine core nanoparticles were used to treat MCF-7 cell spheroids, they effectively targeted the tumor and overcame resistance, angiogenesis, and metastasis. The spheroids' centres were severely damaged [86]. Another therapeutic strategy, using liposomal gold nanoparticles to encase curcumin, was described by Alvi et al. Cur nanoparticles) to treat acne that has returned after developing antibiotic resistance. It used a dual light-mediated treatment with a size of 100-120 nm. By exerting a positive zeta by iontophoresis, these nanoparticles take advantage of targeted follicular delivery [87]. Li and colleagues have effectively created ICG based molybdenum disulfide nanoparticles. With this formulation, the PTT-PDT combinatorial action is achieved, and successfully pglycoprotein, protein is inhibited. By removing anticancer medications from the cell and inhibiting P-glycoprotein, this pump can increase the PDT impact and ultimately lead to Multi Drug Resistance (MDR) [88]. Guo et al. developed a novel method in their investigation by creating a hybrid material from folic acid and chlorin e6 functionalized GO. This combination of PDT and PTT was intended to work in concert.

Interestingly, the mixture demonstrated a fantastic capacity to enter macrophages efficiently. PTT-PDT was found to have a higher combinatorial effect when destroying cancer cells [89]. Using an easy two-step procedure, Liu *et al.* created ICG-encapsulated HA

surface coated with polydopamine nanoparticles. These synthesized nanoparticles display strong PCE and single oxygen radicle generation. Research revealed that cancer cell growth is inhibited by combining PTT and PDT. A novel photo thermal-photodynamic compound for cancer treatment showed great promise when applied to ICG-encapsulated HA surface coated with polydopamine nanoparticles [90].

Effect of light emitting diode (LED) in photo thermal therapy (PTT)

Because LED light sources are less expensive and have fewer side effects, it focuses on LED light sources for photo thermal therapy. For clinical institutions and research labs, LED lighting. However, the newest LED light sources are excellent at reducing the hazards usually associated with lasers while still producing near-infrared light that is both longer-lasting and of higher intensity. The therapeutic applicability of tiny LED light sources is enhanced when they are developed to illuminate a more extensive treatment area than lasers [94].

Effect of mild hypothermia in photo thermal therapy (PTT)

This differs from traditional laser therapy, which requires a long-term and intensive procedure because of its limited ability to reach deep tissue and lower photo thermal conversion efficiency. Increased laser power to achieve effective tumor death while causing some nonspecific damage to the cytoskeleton and vasculature of bystander cells and the host's antitumor immunity. To sensitize cells to mild temperature rise and, ultimately, stop the proliferation of cancer cells, researchers have focused on using PTT and PDT therapeutic modalities at mild temperatures (42–45 °C), either alone or in combination with other strategies that modulate HSP expression, generate ROS, supervise autophagy, and target specific organelles [95–98]. Huang *et al.* transformed cool tumors into hot tumors using a

temperature-sensitive lipid gel depot. Plate number 1 and an antiprogrammed death ligand 1 antibody were added to the centre, and near-infrared radiation was employed to accelerate the infiltration of tumor-infiltrating cells. Optimizing the approach, *in vivo* tests on mice verified the presence of lymphocytes, increased T-cell activity against cancers, and elevation of programmed death ligand 1 in tumor cells [89].

Future perspective

In the upcoming years of research, safety concerns regarding PTT usage as an anticancer treatment will be closely examined and hopefully resolved. Further study will concentrate on how PTT might be combined with other cancer medications to improve therapeutic outcomes and clinical efficiency. Mild hypothermia will be used to decrease the adverse consequences of PTT, and LED lights rather than laser will be encouraged because of their affordability.

CONCLUSION

This review discusses the development of selective biononmaterial's for PTT in cancer treatment. Despite their biocompatibility and biosafety, the long-term toxicity of these materials remains under investigation due to non-biodegradable nanoparticles causing cell toxicity. By optimizing exterior features, bio-nonmaterial's can convert photon energy into thermal energy, enhancing their therapeutic properties. Research aims to identify the best agents for PTT and synergistic therapies like PDT, gene therapy, radio therapy, chemotherapy, and immunotherapy. PTT is beneficial for eradicating primary tumors or lymphatic metastases in superficial tissues. However, the penetration depth of nearinfrared light is limited. To improve cancer treatment, PTT must be integrated with other therapies. Near-infraed-II materials, with higher PCE potential and radiation limits, offer better tissue penetration. Although contrast-enhanced PTT is still under investigation, non-contrast PTT medical devices show promise. Clinical trials continue to explore laser ablation systems without external contrast agents, focusing on practical applications and integration into cancer treatment.

ACKNOWLEDGMENT

The illustrations for their view are prepared using Biorender. com.

FUNDING

Nil

AUTHORS CONTRIBUTIONS

Lokeshvar R: Writing-original draft, Supervision.

 $\label{eq:constraint} \mbox{Yokesh S: Writing-original draft, Methodology, Investigation,} \mbox{Conceptualization.}$

Teejeswari R: Writing-review and editing, Validation.

CONFLICT OF INTERESTS

The authors declare that there are no conflicts of interest in this article.

REFERENCES

- Zhao L, Zhang X, Wang X, Guan X, Zhang W, Ma J. Recent advances in selective photothermal therapy of tumor. J Nanobiotechnology. 2021 Oct 24;19(1):335. doi: 10.1186/s12951-021-01080-3, PMID 34689765, PMCID PMC8543909.
- Tutty MA, Vella G, Vennemann A, Wiemann M, Prina Mello A. Evaluating nanobiomaterial-induced DNA strand breaks using the alkaline comet assay. Drug Deliv Transl Res. 2022 Sep;12(9):2243-58. doi: 10.1007/s13346-022-01178-7. Erratum in: Drug DelivTransl Res. 2022 Jun 19. PMID: 35612707. PMCID PMC9360128.
- Han HS, Choi KY. Advances in nanomaterial-mediated photothermal cancer therapies: toward clinical applications. Biomedicines. 2021 Mar 16;9(3):305. doi:

- $10.3390/biomedicines 9030305, \quad PMID \quad 33809691, \quad PMCID \\ PMC8002224.$
- He CF, Wang SH, Yu YJ, Shen HY, Zhao Y, Gao HL. Advances in biodegradable nanomaterials for photothermal therapy of cancer. Cancer Biol Med. 2016;13(3):299-312. doi: 10.20892/j.issn.2095-3941.2016.0052, PMID 27807498.
- Kim JW, Galanzha EI, Shashkov EV, Moon HM, Zharov VP. Golden carbon nanotubes as multimodal photoacoustic and photothermal high-contrast molecular agents. Nat Nanotechnol. 2009 Oct;4(10):688-94. doi: 10.1038/nnano.2009.231, PMID 19809462, PMCID PMC3663138.
- Liu T, Zhang M, Liu W, Zeng X, Song X, Yang X. Metal ion/tannic acid assembly as a versatile photothermal platform in engineering multimodal nanotheranostics for advanced applications. ACS Nano. 2018 Apr 24;12(4):3917-27. doi: 10.1021/acsnano.8b01456, PMID 29578680.
- He CF, Wang SH, Yu YJ, Shen HY, Zhao Y, Gao HL. Advances in biodegradable nanomaterials for photothermal therapy of cancer. Cancer Biol Med. 2016 Sep;13(3):299-312. doi: 10.20892/j.issn.2095-3941.2016.0052, PMID 27807498, PMCID PMC5069834.
- Xiong X, Wang L, He S, Guan S, Li D, Zhang M. Vacancy defectpromoted nanomaterials for efficient phototherapy and phototherapy-based multimodal synergistic therapy. Front Bioeng Biotechnol. 2022 Aug 25;10:972837. doi: 10.3389/fbioe.2022.972837, PMID 36091444, PMCID PMC9452887.
- Du C, Wu X, He M, Zhang Y, Zhang R, Dong CM. Polymeric photothermal agents for cancer therapy: recent progress and clinical potential. J Mater Chem B. 2021 Feb 19;9(6):1478-90. doi: 10.1039/d0tb02659j, PMID 33427844.
- 10. Fan L, Xu X, Zhu C, Han J, Gao L, Xi J. Tumor catalytic-photothermal therapy with yolk-shell Gold@Carbon nanozymes. ACS Appl Mater Interfaces. 2018 Feb 7;10(5):4502-11. doi: 10.1021/acsami.7b17916, PMID 29341583.
- Yang Y, Zhu D, Liu Y, Jiang B, Jiang W, Yan X. Platinum-carbonintegrated nanozymes for enhanced tumor photodynamic and photothermal therapy. Nanoscale. 2020 Jul 2;12(25):13548-57. doi: 10.1039/d0nr02800b, PMID 32555859.
- Nie D, Ling Y, Lv W, Liu Q, Deng S, Shi J. In situ attached photothermal immunomodulation-enhanced Nanozyme for the inhibition of postoperative malignant glioma recurrence. ACS Nano. 2023 Jul 25;17(14):13885-902. doi: 10.1021/acsnano.3c03696, PMID 37399132.
- Zhang H, Lu XF, Wu ZP, Lou XW. Emerging multifunctional single-atom catalysts/nanozymes. ACS Cent Sci. 2020 Aug 26;6(8):1288-301. doi: 10.1021/acscentsci.0c00512, PMID 32875072, PMCID PMC7453415.
- 14. Zhang Z, Cao Q, Xia Y, Cui C, Qi Y, Zhang Q. Combination of biodegradable hydrogel and antioxidant bioadhesive for treatment of breast cancer recurrence and radiation skin injury. Bioact Mater. 2024;31:408-21. doi: 10.1016/j.bioactmat.2023.08.021, PMID 37692912.
- 15. Melamed JR, Edelstein RS, Day ES. Elucidating the fundamental mechanisms of cell death triggered by photothermal therapy. ACS Nano. 2015 Jan 27;9(1):6-11. doi: 10.1021/acsnano.5b00021, PMID 25590560.
- 16. Manivasagan P, Joe A, Han HW, Thambi T, Selvaraj M, Chidambaram K. Recent advances in multifunctional nanomaterials for photothermal-enhanced fenton-based chemodynamic tumor therapy. Mater Today Bio. 2022;13:100197. doi: 10.1016/j.mtbio.2021.100197, PMID 35036895.
- Wu X, Suo Y, Shi H, Liu R, Wu F, Wang T. Deep-tissue photothermal therapy using laser illumination at NIR-IIa window. Nanomicro Lett. 2020 Jan 24;12(1):38. doi: 10.1007/s40820-020-0378-6, PMID 34138257, PMCID PMC7770864.
- 18. Zhang Y, Zhan X, Xiong J, Peng S, Huang W, Joshi R. Temperature-dependent cell death patterns induced by functionalized gold nanoparticle photothermal therapy in melanoma cells. Sci Rep. 2018 Jun 7;8(1):8720. doi: 10.1038/s41598-018-26978-1, PMID 29880902, PMCID PMC5992202.

- Zhao J, Long X, Zhou M. Clearable nanoparticles for cancer photothermal therapy. Adv Exp Med Biol. 2021;1295:121-34. doi: 10.1007/978-3-030-58174-9 6, PMID 33543458.
- Wang X, Geng Z, Cong H, Shen Y, Yu B. Organic semiconductors for photothermal therapy and photoacoustic imaging. ChemBioChem. 2019 Jul 1;20(13):1628-36. doi: 10.1002/cbic.201800818, PMID 30690811.
- Wang J, Wu X, Shen P, Wang J, Shen Y, Shen Y. Applications of inorganic nanomaterials in photothermal therapy based on combinational cancer treatment. Int J Nanomedicine. 2020 Mar 19;15:1903-14. doi: 10.2147/IJN.S239751, PMID 32256067, PMCID PMC7094149.
- Niu Q, Sun Q, Bai R, Zhang Y, Zhuang Z, Zhang X. Progress of nanomaterials-based photothermal therapy for oral squamous cell carcinoma. Int J Mol Sci. 2022 Sep 9;23(18):10428. doi: 10.3390/ijms231810428, PMID 36142341, PMCID PMC9499573.
- Alamdari SG, Amini M, Jalilzadeh N, Baradaran B, Mohammadzadeh R, Mokhtarzadeh A. Recent advances in nanoparticle-based photothermal therapy for breast cancer. J Control Release. 2022 Sep;349:269-303. doi: 10.1016/j.jconrel.2022.06.050, PMID 35787915.
- Chen Z, Zhuang J, Pang J, Liu Z, Zhang P, Deng H. Application of a cationic amylose derivative loaded with single-walled carbon nanotubes for gene delivery therapy and photothermal therapy of colorectal cancer. J Biomed Mater Res A. 2022 May;110(5):1052-61. doi: 10.1002/jbm.a.37351, PMID 34994069, PMCID PMC9302136.
- Zhang B, Wang H, Shen S, She X, Shi W, Chen J. Fibrin-targeting peptide CREKA-conjugated multi-walled carbon nanotubes for self-amplified photothermal therapy of tumor. Biomaterials. 2016 Feb;79:46-55. doi: 10.1016/j.biomaterials.2015.11.061, PMID 26695116.
- Shinde VR, Khatun S, Thanekar AM, Hak A, Rengan AK. Lipid-coated red fluorescent carbon dots for imaging and synergistic phototherapy in breast cancer. Photodiagnosis Photodyn Ther. 2023;41:103314. doi: 10.1016/j.pdpdt.2023.103314, PMID 36736548.
- Liu R, Zhang H, Zhang F, Wang X, Liu X, Zhang Y. Polydopamine doped reduced graphene oxide/mesoporous silica nanosheets for chemo-photothermal and enhanced photothermal therapy. Mater Sci Eng C Mater Biol Appl. 2019 Mar;96:138-45. doi: 10.1016/j.msec.2018.10.093, PMID 30606519.
- Liang J, Chen B, Hu J, Huang Q, Zhang D, Wan J. pH and thermal dual-responsive graphene oxide nanocomplexes for targeted drug delivery and photothermal-Chemo/Photodynamic synergetic therapy. ACS Appl Bio Mater. 2019 Dec 16;2(12):5859-71. doi: 10.1021/acsabm.9b00835, PMID 35021507.
- Barrera CC, Groot H, Vargas WL, Narvaez DM. Efficacy and molecular effects of a reduced graphene oxide/Fe304 nanocomposite in photothermal therapy against cancer. Int J Nanomedicine. 2020 Aug 25;15:6421-32. doi: 10.2147/IJN.S256760. PMID 32922009, PMCID PMC7457756.
- Zhang YJ, Li BA, Li ZY, Xia N, Yu HY, Zhang YZ. Synthesis and characterization of tamoxifen citrate modified reduced graphene oxide nano sheets for breast cancer therapy. J Photochem Photobiol B. 2018 Mar;180:68-71. doi: 10.1016/j.jphotobiol.2017.12.017, PMID 29413703.
- Gai LX, Wang WQ, Wu X, Su XJ, Yang FC. NIR absorbing reduced graphene oxide for photothermal Radiotherapy for treatment of esophageal cancer. J Photochem Photobiol B. 2019 May;194:188-93. doi: 10.1016/j.jphotobiol.2019.03.014, PMID 31004866
- 32. Liu H, Li C, Qian Y, Hu L, Fang J, Tong W. Magnetic-induced graphene quantum dots for imaging-guided photothermal therapy in the second near-infrared window. Biomaterials. 2020 Feb;232:119700. doi: 10.1016/j.biomaterials.2019.119700, PMID 31881379.
- 33. Qing W, Xing X, Feng D, Chen R, Liu Z. Indocyanine green loaded pH-responsive bortezomib supramolecular hydrogel for synergistic chemo-photothermal/photodynamic colorectal cancer therapy. Photodiagnosis Photodyn Ther.

- 2021;36:102521. doi: 10.1016/j.pdpdt.2021.102521, PMID 34481977.
- Wang Y, Niu C, Fan S, Li Y, Li X, Dai Y. Indocyanine green loaded modified mesoporous silica nanoparticles as an effective photothermal nanoplatform. Int J Mol Sci. 2020 Jul 6;21(13):4789. doi: 10.3390/ijms21134789, PMID 32640753, PMCID PMC7369735.
- Yin Y, Li Y, Wang S, Dong Z, Liang C, Sun J. MSCs-engineered biomimetic PMAA nanomedicines for multiple bioimaging-guided and photothermal-enhanced Radiotherapy of NSCLC. J Nanobiotechnology. 2021 Mar 20;19(1):80. doi: 10.1186/s12951-021-00823-6, PMID 33743720, PMCID PMC7981797.
- 36. Pemmaraju D, Appidi T, Minhas G, Singh SP, Khan N, Pal M. Chlorophyll rich biomolecular fraction of A. Cadamba loaded into polymeric nanosystem coupled with photothermal therapy: a synergistic approach for cancer theranostics. Int J Biol Macromol. 2018 Apr 15;110:383-91. doi: 10.1016/j.ijbiomac.2017.09.084. PMID 28964837.
- Meng F, Han N, Yeo Y. Organic nanoparticle systems for spatiotemporal control of multimodal chemotherapy. Expert Opin Drug Deliv. 2017 Mar;14(3):427-46. doi: 10.1080/17425247.2016.1218464, PMID 27476442, PMCID PMC5385259.
- 38. Huang X, Tang L, Xu L, Zhang Y, Li G, Peng W. A NIR-II light-modulated injectable self-healing hydrogel for synergistic photothermal/chemodynamic/chemo-therapy of melanoma and wound healing promotion. J Mater Chem B. 2022 Oct 5;10(38):7717-31. doi: 10.1039/d2tb00923d, PMID 35920389.
- 39. Wang Y, Yang Y, Yang L, Lin Y, Tian Y, Ni Q. Gold Nanostar@Polyaniline theranostic agent with high photothermal conversion efficiency for photoacoustic imaging-guided anticancer phototherapy at a low dosage. ACS Appl Mater Interfaces. 2022 Jun 29;14(25):28570-80. doi: 10.1021/acsami.2c05679, PMID 35726862.
- 40. Xu M, Zhao D, Chen Y, Chen C, Zhang L, Sun L. Charge reversal polypyrrole nanocomplex-mediated gene delivery and photothermal therapy for effectively treating papillary thyroid cancer and inhibiting lymphatic metastasis. ACS Appl Mater Interfaces. 2022 Mar 30;14(12):14072-86. doi: 10.1021/acsami.1c25179, PMID 35289594.
- 41. Zhang L, Sheng D, Wang D, Yao Y, Yang K, Wang Z. Bioinspired multifunctional melanin-based nanoliposome for photoacoustic/magnetic resonance imaging-guided efficient photothermal ablation of cancer. Theranostics. 2018 Feb 7;8(6):1591-606. doi: 10.7150/thno.22430, PMID 29556343, PMCID PMC5858169.
- 42. Zhang Y, Hong H, Sun B, Carter K, Qin Y, Wei W. Surfactant-stripped naphthalocyanines for multimodal tumor theranostics with upconversion guidance cream. Nanoscale. 2017 Mar 9;9(10):3391-8. doi: 10.1039/c6nr09321c, PMID 28247896, PMCID PMC5435468
- 43. Pan H, Li S, Kan JL, Gong L, Lin C, Liu W. A cruciform phthalocyanine pentad-based NIR-II photothermal agent for highly efficient tumor ablation. Chem Sci. 2019 Jul 18;10(35):8246-52. doi: 10.1039/c9sc02674f, PMID 31673325, PMCID PMC6788506.
- 44. Kumar P, Roy I. Applications of gold nanoparticles in clinical medicine. Int J Pharm Pharm Sci. 2016 Jul;8(7):9-16.
- Duan S, Hu Y, Zhao Y, Tang K, Zhang Z, Liu Z. Nanomaterials for photothermal cancer therapy. RSC Adv. 2023 May 11;13(21):14443-60. doi: 10.1039/d3ra02620e, PMID 37180014, PMCID PMC10172882.
- 46. Liang X, Shang W, Chi C, Zeng C, Wang K, Fang C. Dyeconjugated single-walled carbon nanotubes induce photothermal therapy under the guidance of near-infrared imaging. Cancer Lett. 2016 Dec 28;383(2):243-9. doi: 10.1016/j.canlet.2016.09.006, PMID 27693557.
- Virani NA, Davis C, McKernan P, Hauser P, Hurst RE, Slaton J. Phosphatidylserine targeted single-walled carbon nanotubes for photothermal ablation of bladder cancer. Nanotechnology. 2018 Jan 19;29(3):035101. doi: 10.1088/1361-6528/aa9c0c, PMID 29160225.

- 48. Lu GH, Shang WT, Deng H, Han ZY, Hu M, Liang XY. Targeting carbon nanotubes based on IGF-1R for photothermal therapy of orthotopic pancreatic cancer guided by optical imaging. Biomaterials. 2019 Mar;195:13-22. doi: 10.1016/j.biomaterials.2018.12.025, PMID 30599289.
- McKernan P, Virani NA, Faria GN, Karch CG, Prada Silvy R, Resasco DE. Targeted single-walled carbon nanotubes for photothermal therapy combined with immune checkpoint inhibition for the treatment of metastatic breast cancer. Nanoscale Res Lett. 2021 Jan 7;16(1):9. doi: 10.1186/s11671-020-03459-x, PMID 33411055, PMCID PMC7790975.
- 50. Angoth B, Lingabathula H, Yellu NR. Assessment of extrapulmonary toxicity induced by carbon nanomaterials following intra-tracheal instillation in rats. Asian J Pharm Clin Res. 2017 May;10(5):82-7. doi: 10.22159/ajpcr.2017.v10i5.17152.
- 51. Baez DF. Graphene-based nanomaterials for photothermal therapy in cancer treatment. Pharmaceutics. 2023 Sep 6;15(9):2286. doi: 10.3390/pharmaceutics15092286, PMID 37765255, PMCID PMC10535159.
- 52. Wang Y, Leng S, Huang J, Shu M, Papavassiliou DV. Modeling of cancer photothermal therapy using near-infrared radiation and functionalized graphene nanosheets. Int J Numer Method Biomed Eng. 2020 Jan;36(1):e3275. doi: 10.1002/cnm.3275, PMID 31680480.
- Lu H, Li W, Qiu P, Zhang X, Qin J, Cai Y. MnO2 doped graphene nanosheets for carotid body tumor combination therapy. Nanoscale Adv. 2022 Sep 7;4(20):4304-13. doi: 10.1039/d2na00086e, PMID 36321141, PMCID PMC9552922.
- 54. Qi Y, Qian Z, Yuan W, Li Z. Injectable and self-healing nanocomposite hydrogel loading needle-like nanohydroxyapatite and graphene oxide for synergistic tumour proliferation inhibition and photothermal therapy. J Mater Chem B. 2021 Dec 8;9(47):9734-43. doi: 10.1039/d1tb01753e, PMID 34787633.
- 55. Baipaywad P, Ryu N, Im SS, Lee U, Son HB, Kim WJ. Facile preparation of poly(N-isopropylacrylamide)/graphene oxide nanocomposites for chemo-photothermal therapy. Des Monomers Polym. 2022 Aug 15;25(1):245-53. doi: 10.1080/15685551.2022.2111854, PMID 36017475, PMCID PMC9397426.
- Dash BS, Jose G, Lu YJ, Chen JP. Functionalized reduced graphene oxide as a versatile tool for cancer therapy. Int J Mol Sci. 2021 Mar 15;22(6):2989. doi: 10.3390/ijms22062989, PMID 33804239, PMCID PMC8000837.
- Liu X, Wu X, Xing Y, Zhang Y, Zhang X, Pu Q. Reduced graphene oxide/mesoporous silica nanocarriers for pH-triggered drug release and photothermal therapy. ACS Appl Bio Mater. 2020 May 18;3(5):2577-87. doi: 10.1021/acsabm.9b01108, PMID 35025390.
- Dash BS, Lu YJ, Pejrprim P, Lan YH, Chen JP. Hyaluronic acid-modified, IR780-conjugated and doxorubicin-loaded reduced graphene oxide for targeted cancer chemo/photothermal/photodynamic therapy. Biomater Adv. 2022;136:212764. doi: 10.1016/j.bioadv.2022.212764, PMID 35929292.
- Lima Sousa R, Alves CG, Melo BL, Moreira AF, Mendonça AG, Correia IJ. Poly(2-ethyl-2-oxazoline) functionalized reduced graphene oxide: optimization of the reduction process using dopamine and application in cancer photothermal therapy. Mater Sci Eng C Mater Biol Appl. 2021 Nov;130:112468. doi: 10.1016/j.msec.2021.112468, PMID 34702543.
- Lv S, Miao Y, Liu D, Song F. Recent development of photothermal agents (PTAs) based on small organic molecular dyes. ChemBioChem. 2020 Aug 3;21(15):2098-110. doi: 10.1002/cbic.202000089, PMID 32202062.
- 61. Miranda D, Wan C, Kilian HI, Mabrouk MT, Zhou Y, Jin H. Indocyanine green binds to DOTAP liposomes for enhanced optical properties and tumor photoablation. Biomater Sci. 2019 Aug 1;7(8):3158-64. doi: 10.1039/c9bm00551j, PMID 31232421, PMCID PMC6650340.
- Pebam M, PSR, Gangopadhyay M, Thatikonda S, Rengan AK. Terminalia chebula polyphenol and near-infrared dye-loaded poly(lactic acid) nanoparticles for imaging and photothermal

- therapy of cancer cells. ACS Appl Bio Mater. 2022 Nov 21;5(11):5333-46. doi: 10.1021/acsabm.2c00724, PMID 36288561.
- 63. Jogdand A, Alvi SB, Rajalakshmi PS, Rengan AK. NIR-dye based mucoadhesive nanosystem for photothermal therapy in breast cancer cells. J Photochem Photobiol B. 2020 Jul;208:111901. doi: 10.1016/j.jphotobiol.2020.111901, PMID 32480202.
- 64. Xing R, Liu Y, Zou Q, Yan X. Self-assembled injectable biomolecular hydrogels towards phototherapy. Nanoscale. 2019 Nov 28;11(46):22182-95. doi: 10.1039/c9nr06266a, PMID 31728467.
- 65. Mudigunda SV, Pemmaraju DB, Paradkar S, Puppala ER, Gawali B, Upadhyayula SM et al. Multifunctional polymeric nanoparticles for Chemo/Phototheranostics of retinoblastoma. ACS Biomater Sci Eng. 2022 Jan 10;8(1):151-60. doi: 10.1021/acsbiomaterials.1c01234, PMID 34933546.
- 66. Appidi T, PSR, Chinchulkar SA, Pradhan A, Begum H, Shetty V. A plasmon-enhanced fluorescent gold coated novel lipopolymeric hybrid nanosystem: synthesis, characterization and application for imaging and photothermal therapy of breast cancer. Nanoscale. 2022 Jun 30;14(25):9112-23. doi: 10.1039/d2nr01378a, PMID 35722896.
- 67. Chen J, Ning C, Zhou Z, Yu P, Zhu Y, Tan G. Nanomaterials as photothermal therapeutic agents. Prog Mater Sci. 2019 Jan;99:1-26. doi: 10.1016/j.pmatsci.2018.07.005, PMID 30568319, PMCID PMC6295417.
- 68. Shi Y, Liu M, Deng F, Zeng G, Wan Q, Zhang X. Recent progress and development on polymeric nanomaterials for photothermal therapy: a brief overview. J Mater Chem B. 2017 Jan 14;5(2):194-206. doi: 10.1039/c6tb02249a, PMID 32263539.
- Chen Y, Sun B, Jiang X, Yuan Z, Chen S, Sun P. Double-acceptor conjugated polymers for NIR-II fluorescence imaging and NIR-II photothermal therapy applications. J Mater Chem B. 2021 Jan 28;9(4):1002-8. doi: 10.1039/d0tb02499f, PMID 33399620.
- Beniwal N, Verma A, Putta CL, Rengan AK. Recent trends in bionanomaterials and non-invasive combinatorial approaches of photothermal therapy against cancer. Nanotheranostics. 2024;8(2):219-38. doi: 10.7150/ntno.91356, PMID 38444743.
- Zhang Y, Hong H, Sun B, Carter K, Qin Y, Wei W. Surfactantstripped naphthalocyanines for multimodal tumor theranostics with upconversion guidance cream. Nanoscale. 2017 Mar 9;9(10):3391-8. doi: 10.1039/c6nr09321c, PMID 28247896, PMCID PMC5435468.
- 72. He Y, Wang M, Fu M, Yuan X, Luo Y, Qiao B. Iron(II) phthalocyanine loaded and AS1411 aptamer targeting nanoparticles: a nanocomplex for dual modal imaging and photothermal therapy of breast cancer. Int J Nanomedicine. 2020 Aug 11;15:5927-49. doi: 10.2147/IJN.S254108, PMID 32848397, PMCID PMC7429213.
- Pan H, Li S, Kan JL, Gong L, Lin C, Liu W. A cruciform phthalocyanine pentad-based NIR-II photothermal agent for highly efficient tumor ablation. Chem Sci. 2019 Jul 18;10(35):8246-52. doi: 10.1039/c9sc02674f, PMID 31673325, PMCID PMC6788506.
- 74. Deng X, Shao Z, Zhao Y. Solutions to the drawbacks of photothermal and photodynamic cancer therapy. Adv Sci (Weinh). 2021 Jan 5;8(3):2002504. doi: 10.1002/advs.202002504, PMID 33552860, PMCID PMC7856884.
- Lu H, Wang W, Li X, Zhang M, Cheng X, Sun K. A carrier-free nanoparticle with dual NIR/acid responsiveness by coassembly of enediyne and IR820 for combined PTT/chemotherapy. J Mater Chem B. 2021 May 19;9(19):4056-64. doi: 10.1039/d1tb00279a, PMID 33949615.
- 76. Li X, Bao W, Liu M, Meng J, Wang Z, Sun M. Polymeric micelle-based nanoagents enable phototriggering combined chemotherapy and photothermal therapy with high sensitivity. Biomater Sci. 2022 Sep 27;10(19):5520-34. doi: 10.1039/d2bm00652a, PMID 35924482.
- Cheon YA, Bae JH, Chung BG. Reduced graphene oxide nanosheet for chemo-photothermal therapy. Langmuir. 2016 Mar 22;32(11):2731-6. doi: 10.1021/acs.langmuir.6b00315, PMID 26930106.

- 78. Liu Y, Han YY, Lu S, Wu Y, Li J, Sun X. Injectable hydrogel platform with biodegradable dawson-type polyoxometalate and R848 for combinational photothermal-immunotherapy of cancer. Biomater Sci. 2022 Mar 2;10(5):1257-66. doi: 10.1039/d1bm01835c, PMID 35080214.
- Balakrishnan PB, Sweeney EE, Ramanujam AS, Fernandes R. Photothermal therapies to improve immune checkpoint blockade for cancer. Int J Hyperthermia. 2020 Dec;37(3):34-49. doi: 10.1080/02656736.2020.1797190, PMID 33426992, PMCID PMC7808273.
- Liu K, Hoover AR, Krawic JR, DeVette CI, Sun XH, Hildebrand WH. Antigen presentation and interferon signatures in B cells driven by localized ablative cancer immunotherapy correlate with extended survival. Theranostics. 2022 Jan 1;12(2):639-56. doi: 10.7150/thno.65773, PMID 34976205, PMCID PMC8692917.
- 81. Liu TI, Lu TY, Yang YC, Chang SH, Chen HH, Lu IL. New combination treatment from ROS-Induced sensitized Radiotherapy with nanophototherapeutics to fully eradicate orthotopic breast cancer and inhibit metastasis. Biomaterials. 2020 Oct;257:120229. doi: 10.1016/j.biomaterials.2020.120229, PMID 32738654.
- 82. Gao M, Huang X, Wu Z, Wang L, Yuan S, Du Z. Synthesis of a versatile mitochondria-targeting small molecule for cancer near-infrared fluorescent imaging and radio/photodynamic/photothermal synergistic therapies. Mater Today Bio. 2022;15:100316. doi: 10.1016/j.mtbio.2022.100316, PMID 35721281.
- 83. Jiang Y, Fan M, Yang Z, Liu X, Xu Z, Liu S. Recent advances in nanotechnology approaches for non-viral gene therapy. Biomater Sci. 2022 Dec 6;10(24):6862-92. doi: 10.1039/d2bm01001a, PMID 36222758.
- 84. Odda AH, Cheang TY, Alesary HF, Liu L, Qian X, Ullah N. A multifunctional α-Fe2O3@PEDOT core-shell nanoplatform for gene and photothermal combination anticancer therapy. J Mater Chem B. 2022 Mar 2;10(9):1453-62. doi: 10.1039/d1tb02625a, PMID 35188170.
- Liu Y, Dai X, Yu B, Chen M, Zhao N, Xu FJ. pH-responsive hyaluronic acid-cloaked polycation/gold nanohybrids for tumor-targeted synergistic photothermal/gene therapy. Biomater Sci. 2022 May 17;10(10):2618-27. doi: 10.1039/d2bm00296e, PMID 35412539.
- 86. PSR, Alvi SB, Begum N, Veeresh B, Rengan AK. Self-assembled fluorosome-polydopamine complex for efficient tumor targeting and commingled photodynamic/photothermal therapy of triple-negative breast cancer. Biomacromolecules. 2021;22(9):3926-40. doi: 10.1021/acs.biomac.1c00744. PMID 34383466
- 87. Alvi SB, Appidi T, Deepak BP, Rajalakshmi PS, Minhas G, Singh SP. The "nano to micro" transition of hydrophobic curcumin crystals leading to in situ adjuvant depots for au-liposome nanoparticle mediated enhanced photothermal therapy. Biomater Sci. 2019 Sep 1;7(9):3866-75. doi: 10.1039/c9bm00932a, PMID 31309204.
- 88. Li S, Yang S, Liu C, He J, Li T, Fu C. Enhanced photothermal-photodynamic therapy by indocyanine green and curcuminloaded layered MoS2 hollow spheres via inhibition of P-glycoprotein. Int J Nanomedicine. 2021 Jan 15;16:433-42. doi: 10.2147/IJN.S275938, PMID 33488079, PMCID PMC7815073.
- 89. Guo S, Song Z, Ji DK, Reina G, Fauny JD, Nishina Y. Combined photothermal and photodynamic therapy for cancer treatment using a multifunctional graphene oxide. Pharmaceutics. 2022 Jun 28;14(7):1365. doi: 10.3390/pharmaceutics14071365, PMID 35890259, PMCID PMC9318106.

- Liu X, Xu N, Pu X, Wang J, Liao X, Huang Z. Combined photothermal-photodynamic therapy by indocyanine green loaded polydopamine nanoparticles enhances anti-mammary gland tumor efficacy. J Mater Chem B. 2022 Jun 22;10(24):4605-14. doi: 10.1039/d2tb00565d, PMID 35642529.
- 91. Kamanlı AF, Yıldız MZ, Ozyol E, Deveci Ozkan A, Sozen Kucukkara E, Guney Eskiler G. Investigation of LED-based photodynamic therapy efficiency on breast cancer cells. Lasers Med Sci. 2021 Apr;36(3):563-9. doi: 10.1007/s10103-020-03061-8, PMID 32577931.
- 92. Heiskanen V, Hamblin MR. Photobiomodulation: lasers vs. light emitting diodes? Photochem Photobiol Sci. 2018 Aug 8;17(8):1003-17. doi: 10.1039/c8pp90049c. Erratum in: Photochem Photobiol Sci. 2018 Oct 31;18(1):259, PMID 30044464, PMCID PMC6091542.
- 93. Nikinmaa S, Moilanen N, Sorsa T, Rantala J, Alapulli H, Kotiranta A. Indocyanine green-assisted and LED-light-activated antibacterial photodynamic therapy reduces dental plaque. Dent J (Basel). 2021 May 3;9(5):52. doi: 10.3390/dj9050052, PMID 34063662, PMCID PMC8147628.
- 94. Zhou R, Zhang M, Xi J, Li J, Ma R, Ren L. Gold nanorods-based photothermal therapy: interactions between biostructure, nanomaterial, and near-infrared irradiation. Nanoscale Res Lett. 2022 Jul 26;17(1):68. doi: 10.1186/s11671-022-03706-3, PMID 35882718, PMCID PMC9325935.
- 95. Liu K, Zhang L, Lu H, Wen Y, Bi B, Wang G. Enhanced mild-temperature photothermal therapy by pyroptosis-boosted ATP deprivation with biodegradable nanoformulation. J Nanobiotechnology. 2023 Feb 23;21(1):64. doi: 10.1186/s12951-023-01818-1, PMID 36823540, PMCID PMC9948333.
- Jiang Z, Li T, Cheng H, Zhang F, Yang X, Wang S. Nanomedicine potentiates mild photothermal therapy for tumor ablation.
 Asian J Pharm Sci. 2021 Nov;16(6):738-61. doi: 10.1016/j.ajps.2021.10.001, PMID 35027951, PMCID PMC8739255.
- 97. Wang P, Chen B, Zhan Y, Wang L, Luo J, Xu J. Enhancing the efficiency of mild-temperature photothermal therapy for cancer assisting with various strategies. Pharmaceutics. 2022;14(11). doi: 10.3390/pharmaceutics14112279, PMID 36365098, PMCID PMC9695556.
- 98. Li X, Lovell JF, Yoon J, Chen X. Clinical development and potential of photothermal and photodynamic therapies for cancer. Nat Rev Clin Oncol. 2020 Nov;17(11):657-74. doi: 10.1038/s41571-020-0410-2, PMID 32699309.
- Naser Mohammed S, Mishaal Mohammed A, Al-Rawi KF. Novel combination of multi-walled carbon nanotubes and gold nanocomposite for photothermal therapy in human breast cancer model. Steroids. 2022 Oct;186:109091. doi: 10.1016/j.steroids.2022.109091, PMID 35863403.
- 100. Balou S, Shandilya P, Priye A. Carbon dots for photothermal applications. Front Chem. 2022 Oct 12;10:1023602. doi: 10.3389/fchem.2022.1023602, PMID 36311416, PMCID PMC9597315.
- 101. Cheng X, Liu Y, Zhou H, Leng J, Dai X, Wang D. Cantharidinloaded biomimetic MOF nanoparticle cascade to enhance the fenton reaction based on amplified photothermal therapy. Biomater Sci. 2021 Nov 23;9(23):7862-75. doi: 10.1039/d1bm01396c, PMID 34676840.
- 102. Wadhwa A, Mathura V, Lewis SA. Emerging novel nanopharmaceuticals for drug delivery. Asian J Pharm Clin Res. 2018 Jul 1;11(7):35-42. doi: 10.22159/ajpcr.2018.v11i7.25149.