SYNTHESIS AND CHARACTERIZATION OF THIOLATED GUM KONDAGOGU AND EVALUATION AS MUCOADHESIVE POLYMER

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ABSTRACT

Present work focused on thiolation for enhancing the mucoadhesive potential of Gum kondagogu (GK). Thiolation of GK was done by esterification process with 80 % thioglycolic acid in presence of 7N HCl. Thiolated Gum kondagogu (ThioGK) was determined to possess 1.59 ±0.04 mmol of thiol groups/g of the polymer by Ellman's method. ThioGK was characterized by FTIR, NMR, DSC, XRD, and FE-SEM. The tablets were prepared by direct compression using 75 mg of ThioGK and GK. Tablets containing ThioGK (F1) and GK (F2) were subjected to evaluation of weight variation, hardness and friability and show enhanced disintegration time, swelling behavior, drug release and mucoadhesion. *In vitro* drug release of batch F1 exhibits complete release of drug in 24 hr with zero order release kinetics. Comparative mucoadhesive strength was studied using chicken ileum by texture analyzer and revealed higher mucoadhesion of tablet containing ThioGK. From the above study, ThioGK was suitability exploited as mucoadhesive sustained release matrix tablet.

Keywords: Thiolated gum kondagogu, Mucoadhesive study, Diclofenac sodium, Ellman's method.

INTRODUCTION

Tree gums are natural polymers and are, in recent times receiving consideration as biopolymers since they are non-toxic, inexpensive, simply obtainable, readily improved, environmentally friendly and biocompatible1. In the food and pharmaceutical industry, these natural polymers have a number of applications2. Gum kondagogu (GK) is an important forest produce of Andhra Pradesh, India, which is collected by tribals by tapping from the tree of Cochlospermum gossypium DC (Family: Bixaceae). GK is belonging to substituted rhamnogalacturonans class, which is an anionic polysaccharide. It includes rhamnose, galacturonic acid, glucuronic acid, β-d-galactopyranose, α-d-glucose, β-d-glucose, galactose, arabinose, mannose and fructose with sugar linkage of $(1\rightarrow 2)\beta$ -d-Galp, $(1\rightarrow 6)$ β-d-Gal p, (1→4) β-d-Glc p, 4-O-Me-α-d-Glc p, (1→2) α-l-Rha3. It absorbs a large quantity of water by developing thixotropic gels and in the course of previous studies, for it has been discovered as sustained release matrix tablets4, as emulsifying agent5, as a completely green synthesis of noble metal nanoparticles, for the green synthesis of silver nanoparticles with antibacterial application7, for the mucoadhesive microcapsule

preparation in combination with sodium alginate⁸, Modification of release behavior of gum kondagogu has been executed by carboxylation on gum kondagogu polymeric backbone⁹. GK has been used with other polymers in combination, such as gum olibanum and guar gum as a mucoadhesive polymer since it alone cannot promise residence of a drug delivery system (DDs) at the desired site¹⁰. Till date, several drug delivery systems have been designed using mucoadhesive polymers.

The adhesive property of these natural polymers is the cause of their capability to form noncovalent bonds such as van der Waal's interaction, hydrogen bond and ionic interaction. But, because of such weak interaction, DDs cannot remain at a target site for an extensive period of time^{2,11}. This demands the exploration of novel mucoadhesive polymers. Thiolated polymers have been show to form a class of novel mucoadhesive polymers. Various natural polymers such as karaya gum², chitosan¹²⁻¹⁴, pectin¹⁵, xyloglucan¹⁶, tamarind seed polysaccharide¹, hyaluronic acid¹² and xanthan gum¹³ have been improved by thiol immobilization on polymer to increase their mucoadhesive properties.

In the current study, the chemical modification of gum kondagogu has been carried out by thiolation. The characterization of thiolated gum kondagogu was carried

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Fabrication of efavirenz loaded nano-formulation using quality by design (QbD) based approach: Exploring characterizations and in vivo safety



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Keywords: Efavirenz Nanostructured lipid carrier Quality by design Bioavailability Toxicity

ABSTRACT

Quality by design (QbD) approach was practically applied in fabrication of nanostructured lipid carrier (NLC) encapsulating efavirenz (EF) to ensure the quality in product. Initially, risk factors were categorized based on risk priority number (RPN) using risk identification and assessment tools. A central composite rotatable design (CCRD) was employed to assess the influence of critical process parameter (CPP) (pressure of high pressure homogenizer) and critical material attributes (CMAs) (combination of solid lipid and oil; combination of stabilizers) on responses such as particle size, dispersity and entrapment efficiency. ANOVA was applied to evaluate the data for confirmation of statistical significance (p $\,<\,0.05$). The optimum formulation was decided by setting criteria of responses to achieve desired quality product. This formulation was subsequently lyophilized to evaluate solid state characterization. TEM shows spherical particle shape of NLC. The transformation in amorphous state of NLC from crystalline EF was observed by DSC and PXRD. Lack of molecular interactions and intermolecular hydrogen bonding with lipidic atmosphere revealed by FTIR and ¹HNMR respectively. In vitro drug release 91.21% was obtained at the end of 24 h with Higuchi-matrix mechanism. In vivo pharmacokinetic studies improved relative bioavailability 2.95 fold with lower liver toxicity of EF encapsulated in NLC. In conclusion, QbD based approach clearly proved its usefulness to build quality in product resulting high drug encapsulated potential nanocarrier to enhance bioavailability and confirms safety of EF-NLC with promising acceptable criteria.

1. Introduction

Efavirenz (EF) is a leading drug molecule in the regimen of highly active antiretroviral therapy (HAART) for the treatment of human immunodeficiency virus (HIV). Orally active EF was official by FDA in 1998, belongs to the class of non-nucleoside reverse transcriptase inhibitors (NNRTIs) mostly prescribed to treat HIV-1 infection [1,2]. However, its pharmacokinetic is unpredictable when taken orally. This could be due to poor aqueous solubility, low gastro-intestinal (GI) absorption and rapid first-pass metabolism disappointing *in vivo* pharmacokinetic results [3]. Moreover, it is highly lipophilic (Log P=5.4, intrinsic water solubility = $3-9~\mu g/mL$) drug and categorized in biopharmaceutical classification system (BCS) class II (i.e. poor solubility and high permeability) results in low oral bioavailability of 40-45% [4,5]. The development of hepatotoxicity due to prolong administration is another increasingly important issue limiting the clinical applications [6–8].

Therefore, there is necessitate to develop a strategy which

modulates to improve solubility and bioavailability issues of EF. The few works on such issues have been reported. For example, EF loaded nanoemulsion improved bioavailability with dosage adjustable formulation for HIV therapy [9]. Nanosuspension containing EF prepared and characterized $\beta\text{-cyclodextrin}$ ($\beta\text{-CD}$) based polymeric nanosuspension (PNS) to enhance aqueous solubility and dissolution rate as compared to pure drug [10,11]. The optimized Eudragit E100-Efavirenz loaded polymeric nanoparticles developed to increase in dissolution, drug distribution, and bioavailability, which ultimately implies better control over the therapeutic dosing; and physicochemical evaluation confirmed the formulation stability of nanoparticles [12,13].

SLNs prepared for lymph targeting delivery system to understand chylomicron blocking mechanism approach [14], and NLCs engineered for brain targeted delivery through intranasal route [4]. However, no reports have been addressed on issues of hepatotoxicity which develops on prolong oral administration of EF. Therefore, our prime objective was to develop the EF loaded NLCs to augment biopharmaceutical properties.

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Aloin protects against arsenic trioxide induced myocardial membrane damage and release of inflammatory cytokines

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bstract

Aloin exerts concentration-dependent pro-oxidant and antioxidant effects when tested in vitro. Such duality of effects has not been investigated through in vivo studies on aloin. We evaluated the effects of aloin at doses ranging between 1 and 125 mg/kg against the arsenic trioxide (As₂O₃)-induced cardiotoxicity in mice. As₂O₃ (5 mg/kg/day) was intraperitoneally administrated for 10 days. Aloin was administered through oral gavage at 1, 5, 25, and 125 mg/kg/day. As₂O₃ induced rise in ST height and QT interval in ECG, increased oxidative stress, and depleted the antioxidative defense. As₂O₃ increased inflammatory cytokine concentrations in the heart. Aloin dose dependently inhibited the As₂O₃-induced cardiotoxicity. There was no evidence of increased oxidative stress in the low-dose aloin-treated mice receiving As₂O₃. Our results indicate that aloin possesses cardioprotective potentials and its pro-oxidant effects not evident in vivo at tested doses.





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Heterogeneous surface architectured metal-organic frameworks for cancer therapy, imaging, and biosensing: A state-of-the-art review



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ABSTRACT

With recent progress in inorganic material based nanoplatforms for cancer therapy and imaging, multiple nano vehícles have been developed and evaluated. These recent advancements in material science led to the development of metal organic frameworks (MOFs) and nano MOFs (nMOFs) as the potential and versatile delivery platforms for cancer theranostic, With a vast amount of ongoing research on MOFs, various surface architectured MOFs for with variable properties have been developed and tested. The concept of subcellular targeted therapy of cancer has also been employed using MOFs which demonstrated significantly enhanced anticancer therapy. These MOFs have been developed in a way to provide them stimuliresponsive drug release property which can be utilized for externally guided therapy of cancer. Apart from cellular and subcellular targeted platforms and stimuli-responsive platforms, MOFs have also been explored in the field of bioimaging and biosensing. Multiple types of biosensing platforms based on MOFs and nMOFs have been proposed for biosensing of biomolecules related to cancer for sensing and early detection. The bioimaging probes based on MOFs have been employed for multiple diagnostic platforms. The review gives the recent updates for the abovementioned topics along with the toxicity aspects of MOFs for human use. The review overall gives a detailed overview of research done to date in the field of MOFs based nanoplatforms for cancer theranostics.

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Research Article

Agro-Industrial Waste-Mediated Green Synthesis of Silver Nanoparticles and Evaluation of Its Antibacterial Activity

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DOI: 10.5101/nbe.v12i1.p57-66.

Abstract

The development of immaculate etiquette for the green and rapid synthesis of Ag NPs with a natural reducing agent is the spearhead of the expanding field of nanotechnology. Different scientific fraternity with novel natural reducing agents has been contributing numerous strategies daily. Though there is a submerging of many natural reducing agents, still there are plenty of natural precursors remained to be explored. In this research, we fruitfully attempted the synthesis of silver nanoparticles using agrofood industrial waste *Tamarindus* indica shell-husk extract (TSE) as a natural reducing agent. The prepared silver nanoparticles and their stability in different pH were investigated using ultravioletvisible spectroscopic analysis. Morphological characters were examined using scanning electron microscope (SEM) and transmission electron microscopy (TEM) analysis. The structural and elemental compositions were depicted by Fourier-transform infrared spectroscopy (FTIR) and energy-dispersive X-ray (EDX) analysis, respectively. Moreover, we emphasized on the molecular mechanism involving in the TSE mediated synthesis of Ag NPs. The inherent antimicrobial activity was investigated using agar plate method against both gram-positive and gram-negative species with gentamycin as a control standard for comparison.

Keywords: Green synthesis of Ag NPs; Tamarinds shell-husk extract; Effect of pH; Onepot synthesis; Antimicrobial activity

Introduction

In the recent era, nanobiotechnology has benediction the advantages of the synthesis of nanostructures using living organisms such as plant and microbes. Plant-mediated synthesis of nanoparticles could be advantageous over additional environmentally benevolent biological processes as it eliminates the process involving toxic chemicals and reactants. Biosynthetic processes for nanoparticles would be more expedient if nanoparticles are produced extracellularly using plants or their extracts and in a controlled manner according to their size, dispersity, and shape [1]. Plant-mediated biological synthesis of nanoparticles is gaining significance due to its ease and eco-friendliness. Biosynthetic processes would be more useful if the silver nanoparticles (Ag NPs) were produced using plants or their extracts in a controlled approach according to their dispersity, shape, and size [2]. Although it is a well-known fact

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ORIGINAL ARTICLE



Development of amine-functionalized superparamagnetic iron oxide nanoparticles anchored graphene nanosheets as a possible theranostic agent in cancer metastasis

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Abstract

The major objective of the present investigation was to assess the targeting potential of a designed system for breast cancer at metastatic phases with imaging ability. In a nutshell, we have developed surface-engineered graphene oxide (GO) nanosheets by covalent linking with amine-functionalized iron oxide nanoparticles (IONPs) (GOIOIs). Gefitinib (Gf) was selected as a model drug and entrapped in between exfoliated GO sheets (GOIGF) via π - π * stacking before functionalization with IONPs. Preliminary characterization of GO, IONPs, GOIOI, and GOIGF was performed using UV-visible and Fourier transform infrared spectroscopy. Scanning and transmission electron microscopy studies confirmed successful surface engineering of GO with IONPs. The in vitro drug release study demonstrated sustained release of Gf. The magnetic behavior of IONPs and GOIOI demonstrated a sigmoidal-shaped hysteresis loop with superparamagnetic properties. The in vitro cell cytotoxicity assay was carried out on MDA-MB-231 breast cancer adenocarcinoma cell lines. The cell cytotoxicity assay showed 61.18% inhibition of cell growth with 30 ppm concentration containing 64% of the drug, whereas 100% of the pure drug revealed only 56% of inhibition. In the near future, GOIOI could be tailored further for theranostic research, especially for metastatic cancers.

Keywords Carbodiimide chemistry \cdot Gefitinib \cdot π - π * stacking \cdot MDA-MB-231 breast cancer adenocarcinoma cell lines \cdot Magnetic graphene \cdot Drug delivery

Introduction

Cancer is the most devastating disease in human; one out of six deaths is because of cancer, and the estimated death count may increase up to 13.1 million by 2030. It is the major cause of morbidity and mortality at present. In females, breast cancer is the leading site of cancer followed by cancer of the cervix and uteri [1].

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Chemotherapy [2], gene therapy [3], radiotherapy [4], surgery [5], photodynamic [6], photothermal therapy [7], hyperthermia [8], or a combination thereof have been used for the treatment of cancer. Unfortunately, no effective therapy could successfully eradicate cancer to date. Theranostic nanomedicine is the latest approach under investigation, which could systemically provide simultaneous diagnosis and treatment at a specific site of infection. This could avoid interaction with normal cells, and only cancer tumor cells get destroyed using suitable carrier molecules [9]. The survival rate in cancer patients was dismal from 5 to 15% from developing to developed countries, respectively. Mutation of cancer cell specifically in the epidermal growth factor can be characterized to identify 50% of adenocarcinomas [10].

With the emergence of 2D materials, graphene has gained attention for its use in various biomedical applications including cancer. Graphene is an allotrope of carbon in the form of a single layer of atoms in a two-dimensional hexagonal lattice in which one atom forms each vertex [11, 12]. There are numerous methods available for the synthesis of graphene oxide (GO), an oxidized counterpart of graphene such as mechanical

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Robust Analytical Method for Iron Estimation by Experimental Design Approach

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ABSTRACT

Aim: To perform Iron estimation by UV-Visible spectroscopy using an Experimental design approach. Objectives: The robust analytical method was developed for the estimation of iron (III) using 1, 10-phenanthroline reagent. Methods: The analytical method is an exploration of the chemical reaction of iron with 1, 10- phenanthroline reagent to form a colored complex which was measured in the UV-Visible region at 509 nm. To monitor the effect of diverse factors like the concentration of reagent (A), volume of reagent (B), pH (C) and time (D) on the formation of iron 1, 10 - phenanthroline complex the full factorial design (two-level) was used. From the Pareto chart, Normal plot and half normal plot, it was studied that a combination of all factors was initiate to be significant. Then significant variables are optimized by response surface methodology (RSM) via Box-Behnken design. The evaluation of design was performed to study the effect on the selected response by quadratic effects and main interaction effects. The contour plot and surface plot used for the determined response of the selected factors for their optimum value. Results: The prime reaction state, Beer's law were obeyed in 2.0-10.0 $\mu g/ml$ concentration range with a correlation coefficient of 0.998. Conclusion: The method was successfully applied for the estimation of iron in iron sucrose injection. The optimized method was used for the quantitative analysis of iron sucrose injection.

Key words: Iron sucrose, 1, 10- Pheanthroline, Full factorial design, Box-Behnken design.

INTRODUCTION

Iron is necessary for oxidative metabolism, wound healing, reproduction, cellular growth, execution of several metabolic processes.1 Iron is employed in the production of oxygen-carrying hemoglobin, myoglobin and proteins which are required for the basic metabolic process in the cell.2 Iron deficiency anemia are the most frequent forms of nutritional deficiency generally, anemia is distinct as decrease of hemoglobin value.3 It possesses severe health complications as it causes general weakness, laziness, tiredness, sub-optimal work performance and in certain circumstances psychological obstruction, reduced aptitude and atypical immune response.4 Optimization states to improving the routine of a method, a practice, or produce to get the highest output from it. The term optimization has been generally used in analytical chemistry as a means of

determining situations at which to apply a process that creates the best probable response.⁵

The experimental design is a statistical technique utilized for planning, analyzing and statistical data obtained from primary investigational trials. The experimental design gives exhaustive information from the lowest numeral of trials. Identification of interacting variables characterized the effect of critical factors, evaluation of the effect of preparation and system factors on critical quality attributes.⁶

The conventional optimization approach, varying one variable/factor at a time (OVAT, also called OFAT). One factor at a time (OFAT) does not include interactive outcomes between the variables deliberate as a consequence. OFAT does not include the comprehensive effects of a factor on

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Medicinal Chemistry & Drug Discovery

Exploring Quinazolinones as Anticonvulsants by Molecular Fragmentation Approach: Structural Optimization, Synthesis and Pharmacological Evaluation Studies

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In recent years, design and synthesis of anticonvulsants effective against multiple seizures has attracted much attention of medicinal chemists. In an attempt to find novel anticonvulsants, herein we have reported structurally optimized sixteen different substituted quinazolinones explored through molecular fragmentation approach. The anticonvulsant activity of synthesized compounds was assessed by using predictable seizure models in mice. Two most promising analogues 8 d ($\rm ED_{50}=35.1~mg/kg$, MES, mice, *i.p.*; $\rm ED_{50}=41.5~mg/kg$, scPTZ test, mice, *i.p.*) and 81 ($\rm ED_{50}=21.2~mg/kg$, MES, mice, *i.p.*; $\rm ED_{50}=32.4~mg/kg$, scPTZ test, mice, *i.p.*) exhibited broad spectrum anticonvulsant action in preclinical models of seizures. Com-

pound 81 was also shown profound activity against pharmacoresistant limbic seizures produced in 6 Hz test. Most of the synthesized molecules exhibited moderate to high anticonvulsant activity in all seizure models with no symptoms of neurotoxicity and hepatotoxicity. We have also used *in-silico* protocol for prediction of physiochemical and pharmacokinetic properties of synthesized quinazolinones. The promising anticonvulsant activity of synthesized analogues, *ex-vivo* toxicity, *in-silico* molecular docking, physiochemical and pharmacokinetic predictions make us to anticipate emergence of synthesized quinazolinones as valid leads for the treatment of convulsive disorder.

Introduction

Epilepsy is a grievous and chronic neurological disorder affecting over 65 million individuals worldwide.^[1] An epileptic seizure is abnormal, excessive or synchronous neuronal activity in the brain.^[2] Around 1% of world population at any time is afflicted by epilepsy. This number increases every year by about 2.4 millions.^[3] Antiepileptic drugs (AEDs) therapy is mainstay of treatment for many patients with epilepsy. Development of new AEDs resulted into increased treatment options for patients with epilepsy but making drug selection a more complex task.^[4] Despite decades of research, it is still not clearly understood how AEDs act to control seizures. Unfortunately, at

a present time, there are no AEDs that are significantly effective at different forms and degrees of convulsive disorders.^[5]

Ligand based and random screening protocols have been used by many medicinal chemists to design and synthesize potential anticonvulsants.[6-7] Due to insufficient knowledge and complex pathogenesis of human epilepsy; it is difficult to develop novel AEDs using routine methodologies. Ligand based protocol for design of anticonvulsants is focused on utilization of pharmacophores present in clinically effective and reported AEDs.⁽³⁻⁹⁾ Methaqualone and its synthetic analogues proved as good anticonvulsants. The chloro analogue of methaqualone has shown 1.5 times more potency than phenytoin in MES test and 10 times more efficacy than troxidone in scPTZ model. [6,10-12] Some of the potent anticonvulsants with quinazolinone nucleus are depicted in Figure 1. The substitution of quinazolinone by electron withdrawing halogens at 6th, 7th or 8th positions resulted derivatives with optimum anticonvulsant activity and lower toxicity trian phenytoin.[12-13] Ralitoline is a recently reported AED, found effective in both MES and kindling models of seizures with rodents.[14] Ralitoline has N-(2,6-dimethylphenyl)acetamide as pharmacophoric fragment and shown paramount importance in anticonvulsant potency. Retigabine is an anticonvulsant (ezogabine) approved by the United States Food and Drug Administration (USFDA) and/or by the European Medicines Agency (EMA) in 2011. Retigabine consists of N-(2-aminophenyl)acetamide fragment as critical part of its structure and have vital role in its anticonvulsant activity.[15] Careful incorporation of these pharmacophoric fragments from reported potent anticonvulsants into designed molecules may boost

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PRINCIPAL

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Topical Review

Recent advancements in bioprecursor derived graphene quantum dots: synthesis, characterization and toxicological perspectives

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Abstract

Graphene quantum dots (GQDs), impressive materials with enormous future potential, are reviewed from their inception, including different precursors. Considering the increasing burden of industrial and ecological bio-waste, there is an urgency to develop techniques which will convert biowaste into active moieties of interest. Amongst the various materials explored, we selectively highlight the use of potential carbon containing bioprecursors (e.g. plant-based, amino acids, carbohydrates), and industrial waste and its conversion into GQDs with negligible use of chemicals. This review focuses on the effects of different processing parameters that affect the properties of GQDs, including the surface functionalization, paradigmatic characterization, toxicity and biocompatibility issues of bioprecursor derived GQDs. This review also examines current challenges and the ongoing exploration of potential bioprecursors for ecofriendly GQD synthesis for future applications. This review sheds further light on the electronic and optical properties of GQDs along with the effects of doping on the same. This review may aid in future design approaches and applications of GQDs in the biomedical and materials design fields.

Keywords: bioprecursor, quenching, GQDs, graphene, functionalization of GQDs, hetero-atom doping, fluorescent material

(Some figures may appear in colour only in the online journal)

1. Introduction

With recent advancements in materials sciences and advanced materials, research on the cost of the effective synthesis of materials has gained a lot of attention. Graphene is one of the most celebrated and fascinating 'wonder materials' and is investigated by many branches of science. The graphene family includes graphene, graphene oxide (GO), reduced graphene oxide (rGO) and graphene quantum dots (GQDs). Graphene-based nanomaterials generally exist as

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ORIGINAL ARTICLE



Perspectives of characterization and bioconjugation of gold nanoparticles and their application in lateral flow immunosensing

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Abstract

Gold nanoparticles (AuNPs) are an important component in the field of biomedical diagnostics. Because of its unique physicochemical properties, AuNPs have been widely used in biomedical applications such as photothermal cancer therapy, drug delivery, optical imaging, labeling, and biosensing. In this review, we have described synthesis and characterization techniques for AuNPs with recent advancements. Characterization of AuNPs has played an important role in directing its application in various fields and elaborated understanding of its functioning. The characterization techniques used for the analysis of AuNPs utilize its intrinsic properties, such as surface plasmon resonance (SPR) and size-dependent shift in absorption. These properties of AuNPs are furthermore used for the characterization of bioconjugated AuNPs. Surface conjugation of the AuNPs with biomolecules is explored widely for its use in numerous biosensing applications. Biosensor-based diagnostic devices use AuNPs conjugated with a sensing probe for the detection of a specific analyte. AuNPs are also commonly used as a colorimetric sensor in various point-of-care diagnostic techniques. Lateral flow immunosensing (LFIS) technique utilizes AuNPs for the rapid and sensitive detection of various analytes. LFIS is a paper-based detection technique, where the sample containing the analyte flows through the membrane, interacts with immobilized counterparts, and produces results using a detection probe. AuNPs are used as color markers in LFIS, and the presence of an analyte is indicated by the appearance of colored lines on the membrane. The color is a result of the accumulation of AuNP complexes containing the analyte and probe. Effect of characterization parameters of AuNPs on the sensitivity of LFIS, advantages, and disadvantages of using AuNPs for LFIS are discussed concerning the recent reports. Recent applications of AuNPs in LFIS development for the detection of various biomarkers are summarized comprehensively in the table. The review may offer significant insight into the utility of AuNPs for application in the LFIS technique for future development.

Keywords Gold nanoparticles (AuNPs) · Characterization · Bioconjugation · Lateral flow immunosensing (LFIS) · Diagnostics

Introduction

Nanotechnology is a comparatively new field in research which has been emergent since its introduction as a separate but interdisciplinary subject. Nanomaterial has characteristic properties, which are attributed to its small size and quantum effects. Special physical and chemical properties include high surface to volume ratio, different optical properties from their bulk counterparts, surface plasmon resonance, photothermal effects, fluorescence emission, etc. These unique physical and chemical properties of nanomaterial have enabled their use in a variety of applications such as optical imaging, cancer therapeutics, medical diagnostics, and drug delivery as shown in Fig. 1 [1-9]. The nanomaterial is fabricated using a variety of components, among which metal nanoparticles have gained much importance. Gold nanoparticles (AuNPs) have been proved to be useful for imaging, cancer therapy, and drug delivery [10, 11]. AuNPs have been used for the development of electrochemical immunosensors for detection of Zika virus proteins [12]. Iron nanoparticles, i.e., iron oxide nanoparticles, have been used as nanocarriers to encapsulate anti-HIV drugs

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Central composite design-based optimization of lopinavir vitamin E-TPGS micelle: In vitro characterization and in vivo pharmacokinetic study



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ARTICLE INFO

Keywords: Lopinavir Vitamin E-TPGS Bioavailability enhancement Central composite design

ABSTRACT

This study was aimed at formulating Lopinavir loaded Vitamin E-TPGS micelles to enhance its oral bioavailability. Lopinavir is an HIV-1 protease inhibitor with low aqueous solubility leading to poor oral bioavailability and thus frequent dosing. Drug loaded micelles were fabricated using thin film hydration technique and optimized by two-factor five-level central composite design. For this purpose independent variables selected were TPGS to drug ratio and rotational speed of rotary evaporator, whereas dependent variables chosen were particle size and % entrapment efficiency. The effect of an independent variable on the dependent variable was studied by generating a quadratic polynomial model. Results of in vitro characterization showed that prepared lopinavir micelles exhibited particle size 91.71 nm, polydispersity index 0.129, zeta potential -24.8 mV, entrapment efficiency 99.36 \pm 1.06% and drug loading 20.83 \pm 1.23%. Results of DSC and P-XRD evaluation revealed that drugs were successfully encapsulated inside the Vitamin E-TPGS micelles. In vitro release studies displayed enhancement in drug dissolution as a result of its loading into micelles. TEM images showed that micelles were spherical. On oral administration of lopinavir micelles; the relative bioavailability was boosted by 3.17 folds compared to lopinavir suspensions. Thus, we can conclude that TPGS based micelles possess the prodigious potential to overcome the challenges of current HAART therapy.

1. Introduction

Based on statistics given by the Joint United Nations Programme on HIV and AIDS of the year 2018, nearly 3.79 crores individuals globally are surviving with HIV/AIDS. Out of these 18 lakhs were children of less than 15 years of age. With a rate of 5000 new infections per day about 17 lakhs people universally newly developed HIV infection [1]. Human immunodeficiency virus also abbreviated and commonly known as HIV is a lentivirus. It causes HIV infection which ultimately causes AIDS (Acquired Immunodeficiency Syndrome). AIDS is a condition where the human immune system progressively fails allowing life-threatening opportunistic infections and cancers to conquer the body. It is estimated that normal lifetime post-HIV infection is 9-11 years, based on the HIV subtype [2]. Therefore, an HIV patient needs constant antiretroviral therapy throughout life. This regimen of anti-HIV medication is commonly known as HAART (Highly Active Anti-Retroviral Therapy), which includes a combination of three or more different antiretroviral drugs. Although a complete cure for HIV does not exist, this treatment slows the progression of the virus in the body

by reducing the viral titer in body fluids [3]. Thereby conserving the immune system strength and averting opportunistic infections that may cause death [4].

Lopinavir (LPV) is chemically designated as (2S)-N-[(1S,3S,4S)-1phenoxy)acetyl]amino)-3-hydroxy-5phebenzyl-4-([(2,6-dimethyl nylpentyl]-3-methyl-2-(2-oxotetrahydropyrimidin1 (2H)-yl) butanamide. It is an integral part of the HAART program. But LPV suffers a major drawback i.e., poor bioavailability due to its poor water solubility and cytochrome P450 as well as P-glycoprotein efflux mediated hepatic first-pass metabolism [5]. Thus it is used in combination with ritonavir with the trade names Kaletra® and Aluvia®. Therefore there is a need to develop antiretroviral drug formulation with enhanced bioavailability to improve HAART therapy. Lopinavir and ritonavir both are antiretroviral drugs that are used in combination. Ritonavir is just used as a booster dose for other protease inhibitors and does not have significant antiretroviral activity against HIV and hence it is not prescribed for treatment now a day. It just helps to enhance the bioavailability of Lopinavir. Since after loading into nanocarrier bioavailability of Lopinavir will get enhanced significantly therefore there is no need to use

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Microchemical Journal

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Review Article

A comprehensive review on carbon dots and graphene quantum dots based fluorescent sensor for biothiols



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Keywords:
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ABSTRACT

Fluorescent carbon-based nanomaterials such as carbon dots (CDs) and graphene quantum dots (GQDs) owing to their high aqueous solubility, stable photoluminescence and good biocompatibility are showing greater interest in sensing of biothiols. Biothiols mainly glutathione (GSH), cysteine (Cys), homocysteine (Hcys), considered to be an important tool in the clinical diagnosis of many disorders and diseases. Therefore, the development of new probes has fascinated considerable attention since they are simple, sensitive, rapid and cost effective. Although conventional sensors have been designed and widely applied in biothiols determination, but unfortunately they present many limitations and challenges. In this review, we provide a focused outline on the most recent developments concerning fluorescent based CDs and GQDs nanosensor for detection of biothiols. The most important reaction mechanisms and strategies for detection of biothiols were outlined and compared in terms of their sensitivity and selectivity against different biothiol species and other interfering substances viz. metal ions, amino acids, etc. Future research and challenges in designing of functionalized CDs and GQDs are discussed and elucidated.

1. Introduction

Nanomaterials are the candidates responsible for making breakthroughs in nanotechnology. Over the last two decades, a variety of nanomaterials have been found and evolved as, CDs and GQDs. Quantum effects provide unique features to these nanomaterials when the size of a material reduces to the nanometer range, which failed to predict at macroscopic or microscopic level. As a result, these nanomaterials enhanced the capabilities of researcher to focus on sensing of a variety of materials which could not have been possible with other conventional materials [1]. Nanoparticles are always attributed with the novel properties irrespective of origin [2]. Evolution of the unique physical, chemical and electronic properties at the nanoscale forms the essence of the various applications of nanotechnology [3].

The interplay between the nanomaterials and biological systems for human health concern is of special significance, especially for the CDs and GQDs which have diverse imaging and sensing applications. Biomedical research has become extremely important from past two decades due to human health concern. Biosensing includes qualitative/quantitative recognition of a specific type of analytes by characterizing spectroscopic, electrochemical and photoluminescence behavior of the systems. Most prominently, various biomolecules viz. proteins, nucleic acids, enzymes and chemical analytes e.g., organic metals, inorganic

metals and sugars that help in monitoring the biochemical processes can be detected.

1.1. Fluorescent nanomaterials

Fluorescent nanomaterials viz. CDs, metallic nanoclusters, silicon, metallic nanocomposites, and GQDs immensely revolutionized the field of biosensing and bioimaging [4], however promising CDs [5] and GQDs [6] are attracting increasing attention owing to their high aqueous solubility, low cytotoxicity, stable photoluminescence and better biocompatibility.

1.2. Graphene quantum dots (GQDs)

In recent years, GQDs have gained considerable interest in biosensing and cell imaging applications with other potential applications in diverse areas of the medical and pharmaceutical field [7] owing to their distinctive and remarkable quantum confined electronic state and unique edge structure effect, physicochemical properties [8], fascinating optical properties, high photostability, non-toxicity, biocompatibility and nanometer lateral size [9]. GQDs exhibit excellent photoluminescence properties that can be influenced by structural defect and surface functionality such as heteroatom doping [10].

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Green synthesis of fluorescent graphene quantum dots and its application in selective curcumin detection

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ARTICLEINFO

Keywords: Bamboo timber waste Cellulose nanocrystals Graphene quantum dots (GQDs) Fluorescence resonance energy transfer (FRET) Curcumin detection

ABSTRACT

Herein, we present a facile low-cost and eco-friendly approach for conversion of bamboo timber waste (Bf) derived cellulose nanocrystals (Bf-CNCs) into strong blue luminescent graphene quantum dots (Bf-GQDs) by hydrothermal route. The various properties of synthesized Bf-GQDs were investigated using different spectroscopic techniques. The probable mechanism of Bf-GQDs formation from Bf-CNCs and the effect of pH, particle size on the fluorescent properties of Bf-GQDs also executed. Furthermore, Bf-GQDs were used for the detection of curcumin in an aqueous environment which is the major prerequisite of the present study. The Bf-GQDs showed remarkable photoluminescence (PL) quenching kinetics toward the curcumin (LOD 30.0 nM L⁻¹) assessed by Stern-Volnner plot. The practicability of the method assessed using ginger rhizome juice, while the selectivity of the Bf-GQDs evaluated against different metal ions and different biochemicals. The proposed method will support to establish the strategies for the detection of biochemicals from the aqueous system.

1. Introduction

Graphene based materials, especially graphene quantum dots (GQDs) accolades by researchers amongst the different nanomaterials due to its exceptional properties. Since a scalable GODs production method still stands as a serious obstacle due to its broad application in numerous areas. For example, bottom-up synthesis from graphene and its native precursors demands costly and potentially hazardous chemicals. Moreover, these methods are tedious and required multiple steps to the obtained purified final product (GQDs), which ultimately results in low production yield. Several endeavors have been made worldwide for the conversion of natural waste precursors (NPs) such as weathered leaves [1], coffee grounds [2], etc. Into graphene-like materials and GQDs [3]. It was observed that GQDs from aforementioned precursors has the greatest biocompatibility and stability as compared to the GQDs from pristine graphene as they obtained from a series of hazardous chemical treatment. Moreover, these biomass-derived GQDs are superior to semiconductor quantum dots in terms of solubility, aqueous stability, chemical modification, and high resistance to photo bleaching [4]. These ideal properties have now been subjugated for numerous applications such as fluorescent probes for solar cells [5], biochemical sensing [6] or detection [7,8], bioimaging [9], etc. Apart from the use of available biomaterials such as citric acid [10], sucrose [11], for

obtaining GQDs or graphitic materials one can use waste biomass for gaining one or more active materials of interest. For instance, corn agro waste can be converted into cellulosic derivatives [12], wood waste into quality graphene [13]. The research fraternity well realized that to achieve the 'sustainable green synthesis goals' it is necessary to look after the abundant waste biomass to be utilized for the synthesis of graphene-like materials [3,14,15]. It is well known that cellulose-based biomass hitting the top list as abundant waste at agricultural as well as domestic levels. Cellulose-based polymers had been utilized for the fabrication of numerous materials like nanocomposites in many biomedical and pharmaceutical applications, etc. [16]. A huge amount of bamboo timber waste produces at the various sawmills which further expelled as waste or used as domestic fuel. Bamboo fibers mostly used as a natural composite reinforcing material because of its high strength-to-weight ratio and high hydrophilicity. Cellulose is the major component of the bamboo fibers which plays a crucial role as reinforcing materials in composites while hemicelluloses after the efficient conversion accomplish biofuels and multifunctional by-products. In addition to this hemicellulose on hydrolytic undergoes cleavage of glycosidic bonds between two anhydrous glucose which is the most significant rout of cellulose degradation [17]. Inspired by these facts, we choose the bamboo timber waste for the GQDs synthesis. The stepwise processing of bamboo timber waste yields us cellulose nanocrystals as well as GQDs

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Review

Green Synthesis of Silver Nanoparticles: An Eco-Friendly Approach

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Sopan Namdev Nangare is currently a senior research fellow (Indian Council of Medical Research) at HRPIPER, Shirpur. He has completed his M. Pharm from Bharati Vidyapeeth College of Pharmacy, Kolhapur. He has presented many research papers at various national and international conferences and symposiums. He has been awarded with "Best Outgoing Student" by Bharati Vidyapeeth College of Pharmacy, Kolhapur. He has more than 15 research papers and reviews articles published in national and international journals of repute. His research focuses on the green synthesis of nanocomposites, biosensing, natural polymer-carriers for drug delivery, etc.



Dr. Pravin Onkar Patil received his Ph.D. in January 2014 from R.C. Patel Institute of Pharmaceutical Education & Research, Shirpur (M.S.). He did his M. Pharmacy in Pharmaceutical Chemistry from NDMVPs College of Pharmacy, Nashik (2005). Presently, he is head of the Department of Pharmaceutical Chemistry at HRPIPER, Shirpur, and approved Associate Professor as well as PG teacher in Pharmaceutical Chemistry by North Maharashtra University, Jalgaon. He has presented papers in various conferences, published articles in national and international journals of repute. Recently, he obtained the research grant from SERB (DST), NMU Jalgaon, ICMR, etc. His major fields of scientific interest is green synthesis of graphene-based material for biosensing platform and novel pharmacophore development for several cancer targets using computational tools and their evaluation against a panel of human cancer cell lines using various in vitro assay techniques.

SEAL

Abstract

Eco-friendly synthesis of nanoparticles is an upcoming discipline of nanoscience. Green synthesis of Ag NPs has gained immense importance and much awareness in developed nations. Fascinatingly, such an environmental friendly synthesis of Ag NPs gives a green chemistry-based non-toxic and economical route to nanotechnology. This review article gives insight into the bioinspired synthesis of Ag NPs and mechanisms involved in the synthesis of Ag NPs. In this review, we have summarized the scientific reports in the eco-friendly synthesis arena of Ag NPs and their applications in the biomedical field. Especially, we have focused on plant materials, fungi, algae, and bacterial potential towards the eco-friendly synthesis of Ag NPs. For future perception, there is a need for in silico and in vitro, in vivo research to authenticate the outcomes.

Keywords: Silver nanoparticles, Green synthesis, Plant extract, Eco-friendly, Nanotechnology

http://www.nanobe.org



Silk industry waste protein: isolation, purification and fabrication of electrospun silk protein nanofibers as a possible nanocarrier for floating drug delivery

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Abstract

Amongst assorted regio-selective and targeted oral drug delivery strategies accepted for the gastro-retentive drug delivery system (GRDDS), the floating drug delivery system (FDDS) holds a major share as clinically accepted formulations. The major objective of the present investigation was to explore the silk industry waste protein, silk fibroin (SF) as a possible electrospun nanocarrier for the FDDS. In a nutshell, electrospinning (ES) is one of the flexible and astonishing strategies for the fabrication of porous electrospun nanofibers (NFs), which offers the potential to amend the floating profile, dissolution rate, solubility, and release patterns of the drug, etc as per compendial requirements. Looking at the prospects of floating SF-NFs preparation, we have isolated and lyophilized the SF from industrial waste cocoons and prepared drug-loaded SF single polymer nanofibers (SPN). Lafutidine (LF) being a good candidate for GRDDS selected as a model drug, which is an excellent proton pump inhibitor, mainly used in the treatment of gastric ulcers. Finally, the obtained LF loaded SF-NFs (LF-SF-NFs) were successfully analyzed for physicochemical characteristics, porosity, swelling index, antioxidant activity, mucoadhesion strength, floating properties, enzymatic degradation, and accelerated stability study, etc. Further, these LF-SF-NFs were evaluated for percent drug content, weight variation. in-vitro dissolution in 0.1 N hydrochloric acid (HCl, pH:1.2) and fasted state simulated gastric fluid (FSSGF), and accelerated stability study. It has shown significant floating time > 18 h, about 99% \pm 0.58% floating buoyancy with sustained release up to 24 h. LF-SF-NFs showed good compatibility, entrapment efficiency, antioxidant activity, mucoadhesion strength, enzymatic degradation, and long term stability. Soon, the essential floating and drug release profiles can claim single polymer (SF) based electrospun protein NFs as a possible novel oral nanocarrier for FDDS.

Keywords: processing industrial waste cocoons, silk fibroin, electrospun nanofibers, lafutidine, floating drug delivery system

(Some figures may appear in colour only in the online journal)



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Review

Theranostic Prospects of Graphene Quantum Dots in Breast Cancer

Rahul S. Tade and Pravin O. Patil*



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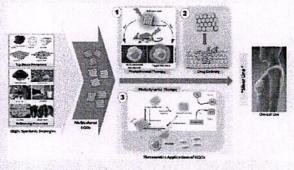
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ABSTRACT: Breast cancer (BC) is increasing as a significant cause of mortality among women. In this context, early diagnosis and treatment strategies for BC are being developed by researchers at the cellular level using advanced nanomaterials. However, immaculate etiquette is the prerequisite for their implementation in clinical practice. Considering the stolid nature of cancer, combining diagnosis and therapy (theranostics) using graphene quantum dots (GQDs) is a prime focus and challenge for researchers. In a nutshell, GQDs is a new shining star among various fluorescent materials, which has acclaimed fame in a short duration in materials science and the biomedical field as well. From this perspective, we review various strategies in BC treatment using GQDs alone or in combination. In addition, the photophysical properties of GQDs



explored in photothermal therapy, hyperthermia therapy, and photodynamic therapy are also discussed. Moreover, we also focus on the strategic use of GQDs both as drug carriers and as combinatorial-guided drug delivery motifs. This Review provides an update for the scientific community to plan and expand advanced theranostic horizons in BC using GQDs.

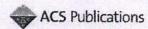
KEYWORDS: breast cancer, triple-negative breast cancer, graphene quantum dots, theranostics, photodynamic/photothermal therapy, drug delivery

1. INTRODUCTION

Cancer is now the second leading cause of death worldwide, with its horrific existence commonly a manner of mortality. In 2018, it was estimated that 9.6 million deaths (i.e., 1 in every 6 deaths) were due to cancer. According to the World Health Organization (WHO), roughly 70% of deaths from cancer occur in low- to middle-income countries (LMICs). Research experts are worried that by the year 2030, 16-18 million additional cases of cancer will be added every year, and 60% of these will be in developing countries. WHO has claimed that by 2030, merely 12 countries are likely to reach a one-third drop in premature cancer mortality.^{2,3} In order to reach sustainable development goals, there is a need for greater investment in the treatment of cancer and other noncommunicable diseases.4 The growing cancer burden globally exerts huge physical, emotional, as well as financial tensions on individuals, families, and consequently the overall health systems. Health organizations in LMICs are least equipped to handle this burden, and for that reason many cancer patients worldwide cannot receive diagnosis and treatment in a timely manner. While the overall cost of cancer treatment in 2010 was estimated at US\$1.16 trillion, only 1 out of every 5 LMICs has the necessary data to drive cancer treatment and mitigation policy.3 Reports suggest that about 30-50% of all cancer cases are predictable, that can be addressed by a cost-effective longterm strategy.5 In many countries, the survival rates of patients diagnosed with cancer are improving owing to prognosis quality treatment and survivorship care.⁶

1.1. Breast Cancer. Every year, breast cancer (BC) affects 2.1 million women, and it is recognized as the most prevalent cancer in women. About 627 000 women died of BC in 2018, which accounts for 15% of all the cancer-associated deaths in women.7 Usually, BC is categorized on the basis of its ability to spread, such as in situ ductal carcinoma (DCIS). DCIS starts in a milk duct and has no further growth into the rest of the breast tissue. Invasive or infiltrating types of BC can spread in the surrounding breast tissue. Invasive BC is comprised of two forms: invasive ductal carcinoma (IDC) and invasive lobular carcinoma (ILC). IDC accounts for 70-80% of all BCs.8 Different forms of BC and global statistics on BC in women are depicted in Figure 1. Invasive BCs have distinct features, which affect their treatment and outlooks. Though invasive BCs are more serious than other types of BC, their occurrence is less common. Invasive BCs are further comprised of inflammatory breast cancer (IBC) and triple-negative breast cancer (TNBC).

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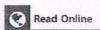
Review

Affinity-Based Nanoarchitectured Biotransducer for Sensitivity Enhancement of Surface Plasmon Resonance Sensors for *In Vitro* Diagnosis: A Review

Sopan N. Nangare and Pravin O. Patil*



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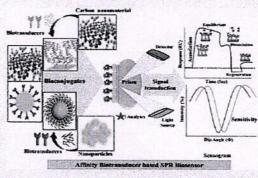


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ABSTRACT: Despite the indisputable benefits and advancement of science, technology, and civilization, early diagnosis of healthcare is still a challenging field for the scientific fraternity. The detection of biomarkers is a crucial attribute of prognosis and diagnosis of disease. Out of numerous techniques, surface plasmon resonance (SPR) bestows countless benefits, including in situ, label-free, and real-time assessment, etc., which authorizes the analysis of molecular binding occurrences between biotransducers and biomarkers. In addition, SPR with low-molecular-weight biomarkers lacks selectivity and sensitivity, which ultimately affects binding kinetics. This, in turn, leads to the remarkable development and implementation of numerous selectivity and sensitivity enhancement methods. Among the various noticeable strategies, because of selectivity and sensitivity enrichment substrate for SPR biosensors, affinity-based nanoarchitectured



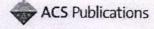
biotransducers stand out as being the best substitute. The present review elaborates significant advances made in the research based on affinity biotransducers for *in vitro* diagnosis using SPR biosensors for biomarker sensing. Moreover, most recent trends and challenges in designing and application of nanoarchitectured affinity biotransducer-based SPR biosensors for detecting low-concentration biomarkers have been reviewed comprehensively. This present review may assist the scientific fraternity in designing an ultramodern novel SPR approach based on affinity biotransducers, along with improved selectivity and sensitivity of SPR biosensors for *in vitro* and real-time diagnostic applications.

KEYWORDS: Affinity biotransducer, surface plasmon sensor, in vitro diagnosis, antibody, aptamer, nanoparticles

1. INTRODUCTION

In addition to the indisputable benefits of research, industry, and technology, unfortunately, some impetuous shifts in the natural world have started to endanger the lives of peoples and other entities directly.1 In the last couple of decades, a key feature to achieving rapid diagnosis of much pathology is the insistence for early, economical, and reliable analytical instruments for in vitro diagnosis.² Initially, the scientific fraternity has fixed the general criterion for in vitro diagnostic devices, which includes the utilization condition and risk factors of the device/machine. In addition, the device should be proficient to provide relevant information for careful diagnosis of particular health issues. Furthermore, the result of analytical techniques or devices should notably affect the public or individual negatively or positively.3 The literature survey revealed that the emerging in vitro diagnostic devices are offering rapid screening and early detection ability, precise information, and real-time monitoring of several diseases and disorders.4 Unfortunately, official analytical strategies (commonly employed for diagnostic applications) are plagued by numerous drawbacks, viz, selectivity, sensitivity, time-consuming process, the cost of analysis, need of expert and trained staff for its laborious process, and having limited availability as a point of health care system, etc. Moreover, the luminous technological expansion within assorted sectors and fields (viz, nanotechnology, biotechnology, and electronics) necessitates impetuous, user-friendly, sensitive tools, which leads to a tremendous expansion of analytical methods in the last decades. More precisely, the biosensor is an advanced analytical tool in which a biotransducer (example: antibodies, aptamers, tissues, DNA, enzymes, etc.) is united through a physicochemical transducer (viz, optical, magnetic, electrochemical, piezoelectric). The interaction between the interest/target biomarker (or analyte) and the specific

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Journal Pre-proof

One- Pot Development of Spray Dried Cationic Proliposomal Dry Powder Insufflation: Optimization, Characterization and Bio-interactions

Ajjappla Basavaraj Shreya, Abhijeet Pandey, Ajinkya Nitin Nikam, Pravin O. Patil, Raju Sonawane, Prashant Deshmukh, Srinivas Mutalik



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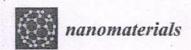
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Review

Black Phosphorus as Multifaceted Advanced Material Nanoplatforms for Potential Biomedical Applications

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Abstract: Black phosphorus is one of the emerging members of two-dimensional (2D) materials which has recently entered the biomedical field. Its anisotropic properties and infrared bandgap have enabled researchers to discover its applicability in several fields including optoelectronics, 3D printing, bioimaging, and others. Characterization techniques such as Raman spectroscopy have revealed the structural information of Black phosphorus (BP) along with its fundamental properties, such as the behavior of its photons and electrons. The present review provides an overview of synthetic approaches and properties of BP, in addition to a detailed discussion about various types of surface modifications available for overcoming the stability-related drawbacks and for imparting targeting ability to synthesized nanoplatforms. The review further gives an overview of multiple characterization techniques such as spectroscopic, thermal, optical, and electron microscopic techniques for providing an insight into its fundamental properties. These characterization techniques are not only important for the analysis of the synthesized BP but also play a vital role in assessing the doping as well as the structural integrity of BP-based nanocomposites. The potential role of BP and BP-based nanocomposites for biomedical applications specifically, in the fields of drug delivery, 3D printing, and wound dressing, have been discussed in detail to provide an insight into the multifunctional role of BP-based nanoplatforms for the management of various diseases, including cancer therapy. The review further sheds light on the role of BP-based 2D platforms such as BP nanosheets along with BP-based 0D platforms—i.e., BP quantum dots in the field of therapy and bioimaging of cancer using techniques such as photoacoustic imaging and fluorescence imaging. Although the review inculcates the multimodal therapeutic as well as imaging role of BP, there is still research going on in this field which will help in the development of BP-based theranostic platforms not only for cancer therapy, but various other diseases.

Keywords: bioimaging; wound healing; 3D printing; surface modification; characterization

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1. Introduction

The discovery of Black Phosphorus (BP) dates back to a hundred years ago. It all began with Bridgman [1], who brought about the conversion of white phosphorus to black phosphorus under a high temperature and pressure. Later, Hultgren et al. [2] demonstrated

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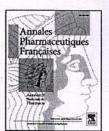
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GENERAL REVIEW

Carbon dots: A novel trend in pharmaceutical applications



Carbon dots: une nouvelle tendance dans les applications pharmaceutiques

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HIGHLIGHTS

- Presents basic concepts, advantages, synthesis approach of CDs.
- Numerous CDs based pharmaceutical applications of were reviewed.
- CDs were used in gene therapy and nanomedicine.
- CDs were used in bioimaging and biosensing.

KEYWORDS

Carbon dots; Pharmaceutical applications; Bioimaging; Sustained drug delivery; Targeted drug delivery Summary Carbon quantum dots (CQDs, C-dots, or CDs), are generally small carbon nanoparticles having a size less than 10 nm. Carbon dots (CDs) were accidentally discovered during the purification of single-walled carbon nanotubes through preparative electrophoresis in 2004. Carbon is an organic material having poor water solubility that emits less fluorescence. However, CDs have good aqueous solubility and excellent fluorescent property, hence more attention has been given to the synthesis of CDs and their applications in chemistry and allied sciences. CDs being easily accessible for in-house synthesis, simpler fabrication as per compendial requirements are wisely accepted. In addition, since CDs are biocompatible, of low toxicity, and of biodegradable nature, they appear as a promising tool for the health care sector. Furthermore, owing to their capabilities of expressing significant interaction with biological materials, and their excellent photoluminescence (PL), CDs have been emerging as novel pioneered nanoparticles useful for pharmaceutical and theranostic applications. Also, CDs are more eco-friendly in synthesis and therefore can be favorably consumed as alternatives in the further development

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