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## Review

## Nanomaterial application in bio/sensors for the detection of infectious diseases

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## ABSTRACT

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Infectious diseases are a potential risk for public health and the global economy. Fast and accurate detection of the pathogens that cause these infections is important to avoid the transmission of the diseases. Conventional methods for the detection of these microorganisms are time-consuming, costly, and not applicable for on-site monitoring. Biosensors can provide a fast, reliable, and point of care diagnostic. Nanomaterials, due to their outstanding electrical, chemical, and optical features, have become key players in the area of biosensors. This review will cover different nanomaterials that employed in electrochemical, optical, and instrumental biosensors for infectious disease diagnosis and how these contributed to enhancing the sensitivity and rapidity of the various sensing platforms. Examples of nanomaterial synthesis methods as well as a comprehensive description of their properties are explained. Moreover, when available, comparative data, in the presence and absence of the nanomaterials, have been reported to further highlight how the usage of nanomaterials enhances the performances of the sensor.

## 1. Introduction

Infectious or transmissible diseases are those diseases that can be transmitted by bacteria, viruses, fungus, and parasites [1]. According to the World Health Organization reports consumption of unsafe foods, that are polluted with harmful bacteria, viruses, parasites, or chemical substances leads to 200 types of sickness including cancers, sepias, homoerotic diarrhea, etc. [2]. Moreover, diarrhea-associated infections related to the intake of contaminated food or water are the third cause of mortality in developing countries, and around a million people's death is reported for this every year [2–4].

Facing threats like Ebola, dengue, Zika, severe acute respiratory syndromes (SARS-CoV-2), Middle East respiratory syndrome (MERS), and severe known as well as unknown pathogens endanger human health and, in a large view, economic welfares [5]. Bacteria such as *Escherichia coli* O157: H7, *Salmonella enterica* serovar *Typhimurium*, *Klebsiella pneumonia*, *Listeria monocytogenes*, *Enterococcus faecalis*, *Campylobacter*, *Acinetobacter baumannii* and Viruses like *influenza virus H5N1*, *H3N2*, *HIV*, *HPV*, and *enterovirus 71* cause enormous

hospitalizations and death all over the world. Therefore, the development of new, fast, and reliable approaches to detect these sources of infection are crucial to prevent the epidemic of these diseases [6,7].

Nanomaterials with specific electrical, optical, magnetic, chemical and mechanical properties have been found effective application in many areas including biosensors or chemical sensors, diagnostics methods, drug delivery, energy harvesting, and tissue engineering [8]. This unquestionably owes to the intrinsic high surface area to volume ratio of these materials that can present more recognition sites and quicker signal transduction procedure [9]. Nanomaterials according to the European commission's recommendation 2011, "is a natural, incidental or manufactured material containing particles, in an unbound state or as an aggregate or as an agglomerate and where, for 50% or more of the particles in the number size distribution, one or more external dimensions is in the size range 1 nm–100 nm" [10]. They can be categorized into four groups based on the material using in their fabrication 1) carbon-based nanomaterials that are carbons with different morphology such as hollow tubes, ellipsoids or spheres e.g. Graphene, Graphene oxide, Carbon nanotubes (CNTs), Fullerenes (C60), Carbon

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black and carbon nano onion. 2) Inorganic-based nanomaterials, these groups involve metal (Au, Ag, Pt, Ni, Cu), metal oxide nanoparticles ( $ZnO$ ,  $TiO_2$ ,  $MnO_2$ ,  $Fe_2O_3$ ,  $Al_2O_3$ ),  $SiO_2$ , semiconductors such as silicon and ceramics, and inorganic semiconductor nanocrystals which also called quantum dots ( $CdS$ ,  $ZnO$ ,  $CdS$ ). 3) Organic-based nanomaterials, these groups are nanomaterial from organic material such as (dendrimers, micelles, liposomes, and polymer NPs). 4) Composite-based nanomaterials, that are multi-phase nanoparticle and nanostructured materials with one phase on the nanoscale size which can combine either with other nanoparticles or bulk-type materials (e.g., hybrid nanofibers) or more complicated structures, such as metal-organic frameworks (MOF). The composites may be any combinations of carbon-based, metal-based, or organic-based nanostructured materials with any form of metal, ceramic, or polymer bulk materials [11,12]. In the present review, we will focus on electrochemical and optical sensors for infectious diseases which used nanomaterials like metallic nanoparticle (Ag, Au), magnetic nanoparticle, bimetallic nanoparticle, silica nanoparticle, graphene family, carbon nanotube, carbon dot, quantum dots, and upconversion nanoparticles.

## 2. Nanomaterial

### 2.1. Magnetic nanoparticle (MNP)

Magnetic nanoparticles are well known for special magnetic properties like superparamagnetism and high magnetic susceptibility [13]. These nanoparticles have found extensive applications due to their unique advantages including low production cost, large surface area, high mass transference, direct capture, easy separation, improving the washing steps, and minimizing the matrix effect. Besides their application in sample enrichment, they have been shown to enhance sensor sensitivity, increase the signal-to-noise ratio, and decrease the time of analysis [14,15]. The role of the MNPs is dependent on their sizes that can be influenced by the preparation method [16]. MNPs with a size of less than 30 nm possess superparamagnetic properties making these good candidates for enrichment applications because of an immediate response to the applied magnetic field [17]. Different types of Magnetic nanoparticles like iron oxides ( $Fe_2O_3$  and  $Fe_3O_4$ ); ferrites of manganese, cobalt, nickel, and magnesium; multifunctional composite MNPs, such as  $Fe_3O_4$ -Ag,  $Fe_3O_4$ -Au,  $Fe_3O_4$ - $SiO_2$  are reported. Among them,  $Fe_3O_4$  is the preferred magnetic nanoparticle in biosensor development because of its superparamagnetic property, biocompatibility with antibodies and enzymes, and ease of preparation [14]. Among the demonstrated manufacturing methods, like thermal decomposition, microemulsion, electrochemical, solvothermal, sol-gel, sonochemical, and co-precipitation for synthesis  $Fe_3O_4$  nanoparticle; co-precipitation is the most used technique in the synthesis  $Fe_3O_4$  [17]. More details about different synthesis method of iron oxide nanoparticles and surface modification [17–19] properties of iron oxide nanoparticles [17], magnetic  $CoFe_2O_4$  [20] magnetic nanoparticle [21] are available in these references.

### 2.2. Gold nanomaterials

Gold nanomaterials with a wide variety of shapes, such as rods, spheres, cubes, wires, and cages, are among the most popular nanomaterials for sensor applications [22]. Their intrinsic properties, such as good biocompatibility, large surface area, ease of synthesis and modification, and high chemical and thermal stability make Au nanomaterial ideal choice as substrates and transducers for sensors. The optical properties of these materials depend on their size and shape which can be controlled/tuned during the synthesis process. Besides, the surface of gold nanomaterial can be modified by electrostatic, hydrophobic, and covalent bonding; for example, the strong bond between gold atoms and nitrogen or sulfur, enable efficient attachment of molecules via the well-established thiol or amine chemistry [9].

### 2.2.1. Gold nanoparticle (AuNPs)

Gold nanoparticles (AuNPs) are polycrystalline Au nanostructure with a quasi-spherical shape and typically 5–100 nm diameter size. In electrochemical sensors, AuNPs, have found application due to their ability to improve the charge transfer processes [23] and the electrode's conductivity [24]. AuNPs are extensively applied as a colorimetric indicator, because of their surface plasmon resonance property [25,26]. Along with their optical and electrochemical properties, AuNPs are very attractive in biosensing since their high surface area offer unprecedented opportunities for high loading of biomolecules such as antibodies, enzymes, and aptamers; moreover, gold nanoparticles have been shown to maintain high bioactivity of immobilized biomolecules [23]. Chemical reduction or ligand passivation methods and their variants are popular approaches for manufacturing AuNPs. Peng et al. discussed various methods for the synthesis of gold nanomaterial family [9].

### 2.2.2. Gold nanorods (AuNRs)

Anisotropic Nanorods are cylindrical rod-shaped particles with uniform diameters ranging from a few nanometers to hundreds of nanometers, with aspect ratios (length to width) larger than 1 but typically smaller than 10 [27]. The main features of AuNRs are depending properties like intensities and spectral positions of SPR-bands, aggregative stability, electroconductivity, and redox potential by physical size [28]. One of the peculiarities of the AuNRs is those to present two distinct surface plasmon resonances; one along with the longitudinal ax and one on the transversal ax. The longitudinal plasmon resonance generates radiation in a wide range from visible to near-infrared regions depending on the AuNRs aspect ratio and their size which is a distinguishable feature of nanorods when compared to nanoparticles [28,29]. Among the different approaches proposed to producing AuNRs, seed-mediated growth is the most commonly used because of a simple procedure, adjustable aspect ratio, great uniformity, and adaptability for post-modification [9,30]. There are promising reviews about the synthesis and properties of gold nanorods [30] gold nanorods and their nanocomposites synthesis and recent applications in Analytical chemistry [28].

### 2.2.3. Gold nanocluster (AuNCs)

AuNCs are molecular like arrangements consisting of several hundred Au atoms with an average size of 2 nm. They are located between Au atoms and AuNPs and can be fabricated by atoms to clusters and nanoparticles to clusters approaches. Atoms to cluster approach based on template-based synthesis and ligand-protected methods. While, nanoparticles to clusters methods are etching surface atoms of gold nanoparticles by appropriate ligand, or solvents [31]. AuNCs are characterized by photoluminescence [32] and high catalytic properties [33]; on the other end, AuNCs do not present the surface plasmon resonance properties characteristic of the AuNPs and AuNRs [9]. Zhang et al. discussed synthesis methods and biomedical and biosensing applications of Au nanoclusters in a recent review [34].

### 2.3. Silver nanoparticles (AgNPs)

Silver nanoparticles (AgNPs) present unique plasmonic properties that can be influenced by different parameters, such as size, shape, uniformity, composition, dispersion, etc. [35]. Moreover, these nanomaterials have prominent benefits including low cost, simple preparation, high extinction coefficients, and sharp extinction band [36]. AgNPs can be produced, similarly to their counterparts AuNPs, by chemical reduction using reducing agents like  $NaBH_4$ , tri-sodium citrate, thiosulfate, and polyethylene glycol. Polyvinyl pyrrolidone (PVP), polyvinyl alcohol (PVA), and CTAB are often used to stabilize particles and avoid sedimentation and agglomeration of nanoparticles [37].

## 2.4. Bimetallic nanoparticles

Bimetallic nanoparticles are consist of two different metal NPs and can have various morphologies and structures. They can be classified based on structures into mixed and segregated structures. Mixed forms subdivide into alloy and intermetallic. In alloyed nanoparticles nanocrystals of two metals are randomly mixed; however, in intermetallic structures, the two metals are mixed in an ordered way. The segregated structures are categorized as subcluster, core-shell, multishell core-shell structures, and multiple core structures. A subcluster is a separate distribution of two metals with a shared interference. Core-shell structures are composed of a metal core surrounded by a shell of the second metal. Multishell core-shell structures are an alternative arrangement of metals forming a shape like onion rings. Multiple core materials are multiple small cores coated by a single shell. Various types of bimetallic nanoparticles are platinum, nickel, iron, palladium, and gold-based bimetallic nanoparticles [38]. Due to the collaborative effect of two different metals, bimetallic nanoparticles display novel features such as tunable surface plasmon band and optical properties and improved stability and dispersibility that make them attractive materials in electrochemical and optical sensors [39,40]. They can prepare with chemical, physical, and biological methods [41]. Some interesting reviews for synthesis, properties, and application of bimetallic nanoparticle [39,42], noble metal bimetallic nanoparticle [43] are provided in the former references.

## 2.5. Quantum dots (QDs)

Quantum dots (QDs) is a family of nanomaterials with outstanding optical properties like narrow emission, wide absorption spectra, size-controlled emission from visible to the infrared range (400–4000 nm), good photostability, and high quantum yields. Generally, these are composed of atoms from IIB-VI, III-V, or IV-VI groups of the periodic table that have 1–10 nm size dimensions which is smaller than the exciton Bohr radius [44–46]; their nanocrystal (dots) nature defines the quantum controlled mechanics of their optical properties. Capping agents on the surface of QD including alcohols, primary amines, carboxylic acid thiols, and long-chain organophosphorates play a pivotal role in the application of these materials. Ligands facilitate modification of the surface with biomolecules and responsible for colloidal stability, solubility, particle size distribution as well as growth or agglomeration [47,48].

## 2.6. Graphene group

Graphene is one of the carbon allotropes with a plane two-dimensional nanostructure. This possesses excellent optical and electronic properties, converting it into a fast-growing material for designing optical and electrochemical sensors [49,50]. There are different graphene-based nanomaterials, pristine graphene, graphene oxide (GO), and reduced graphene oxide (rGO). GO was regarded as a result of chemical exfoliation and oxidizing of layered crystalline graphite [51]. rGO is the product of GO reduction via physical, chemical, or electrochemical methods. These differ from each other in terms of, composition, purity, lateral dimensions, oxygen content, surface chemistry, and electric properties [52,53]. GO is strongly hydrophilic with oxygenated functional groups that facilitate chemical functionalization of it. The presence of versatile functional groups such as OH, COOH, and CHO allows interaction with biomolecules to improve the selectivity of the biosensors [54] but reduce the electrical conductivity due to its insulating nature for electrochemical application [55]. rGO demonstrated an improvement in electrical conductivity and remaining oxygen groups, converted it to a useful material in developing biosensors. Graphene and graphene derivatives can be coupled with different materials, such as metal oxides, metal nanoparticle, and organic polymers obtaining in this way diverse nanocomposites with

tailored properties [56,57]. Interesting reviews about graphene-based electrochemical biosensors that discussed different types and synthesis methods [55] and reduced graphene for electroanalytical sensing platforms [58] are available in the former papers.

## 2.7. Carbon nanotube (CNTs)

Carbon nanotubes (CNTs), including single-wall and multiwall carbon nanotubes (SWCNTs and MWCNTs, respectively) [59] have attracted great attention in sensors due to their high surface area, high electrical conductivity, high thermal conductivity, and great mechanical strength. CNTs also possess specific inherent optical properties as, for example, potent resonance Raman scattering and Near-Infrared photoluminescence [60]. CNT can produce by three main synthesis methods including chemical vapor deposition, electric arc method, and laser deposition method [61].

## 2.8. Carbon dot or carbon particles (CD)

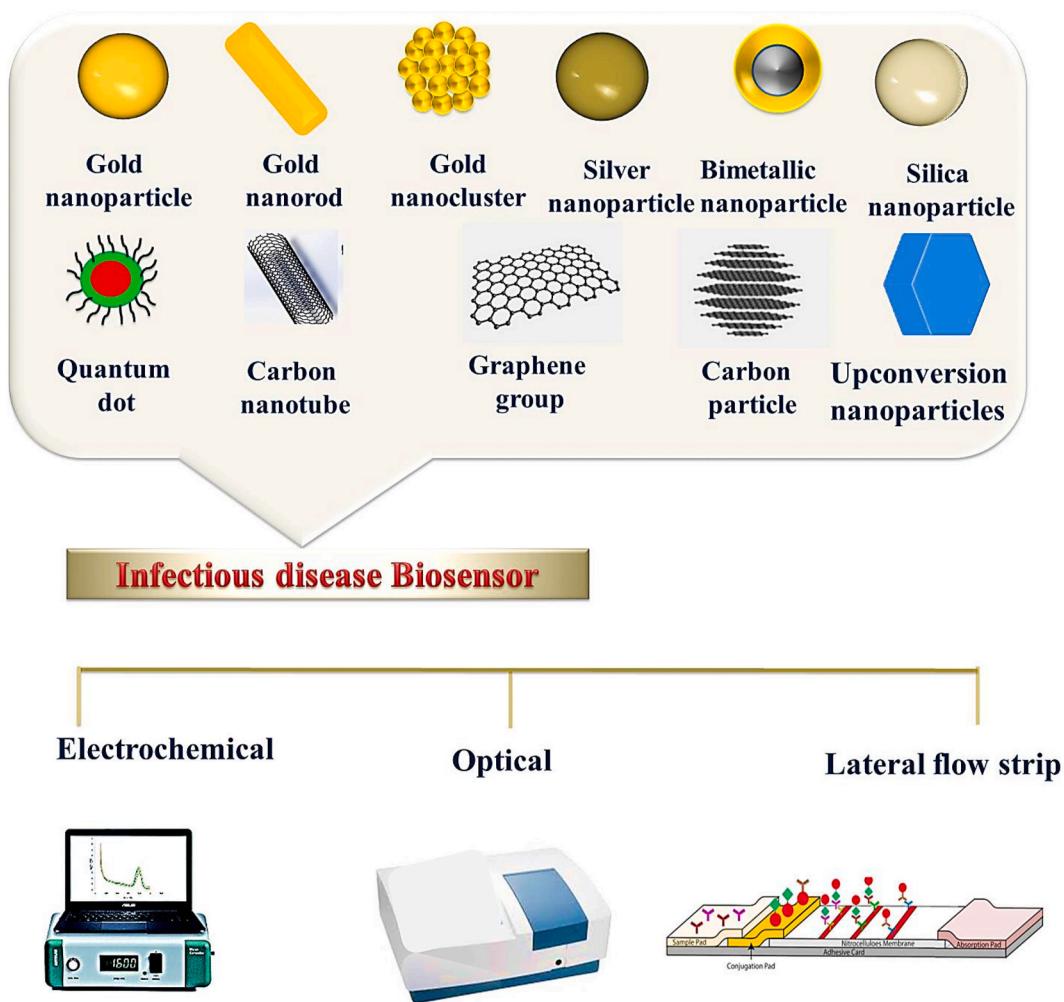
Carbon dots (CDs), as a new type of fluorescent carbon-based nanomaterial, have promising properties to design the fluorescent sensor to conventional fluorescent probes like superior optical properties, strong absorption, bright photoluminescence, excellent light stability, resistance to light bleaching, low toxicity, environmental-friendly, good biocompatibility, and facile synthesis [62,63]. CDs are zero-dimensional carbon-dominated nanomaterial, with a size of less than 20 nm which consisted of  $sp^2/sp^3$  carbon skeleton and abundant functional groups/polymer chains [62]. Functional groups are OH, COOH, and NH<sub>2</sub> [64,65]. Although there is considerable debate about the classification of CDs, they can be categorized into carbon quantum dots (CQDs) graphene quantum dots (GQDs), carbon nanodots (CNDs), and polymer dots (PDs) [62]. Two main synthesis routes, the top-down include arc discharge, laser ablation, electrochemical synthesis, nanometer etching, hydrothermal/solvothermal/special oxidation cleavage, and the bottom-up method involves combustion, pyrolysis, hydrothermal, solvothermal, and microwave assisted pyrolysis have been reported for the preparation of carbon dots [62,63,66].

## 2.9. Upconversion nanoparticles (UCNPs)

Upconversion nanoparticles (UCNPs) with near-infrared (NIR) excitation characteristics have received considerable attention in bio-imaging (e.g. nanoprobe) [67]. The most used of UCNPs-based nanoparticles are in fluorescence resonance energy transfer, FRET sensor. The UCNPs fluorescence were quenched or recovered after the additions of analytes via modulating the absorption of energy acceptors or the distance between donors and acceptors [68]. UCNPs are a special class of nanomaterial doped with lanthanide ions; these consist of three components, a sensitizer ( $Yb^{3+}$ ) for absorbing energy from a laser source, an emitter that works as a luminescence hearts of UCNPs ( $Tm^{3+}$ ,  $Er^{+3}$ ,  $Ho^{+3}$ ), and a host matrix. The host matrix has a substantial influence on optical features, fluorescence efficiency, and chemical properties. Fluorides due to their high chemical stability, low photon energy, are well-suited hosts for UCNPs. Synthetic procedures like co-precipitation, thermal decomposition, hydro/solvothermal synthesis, and sonochemical methods have been reported in the manufacturing of UCNPs [67]. These nanoparticles are characterized by low toxicity, sharp emission band, long fluorescence lifetime, less light scattering, high photostability, tunable multicolor emission, high light penetration depth, and larger anti-stocks shifts, when compared with more conventional fluorochromes, making them particularly useful for biosensor applications [68,69].

## 2.10. Silica nanoparticles

Numerous silica nanoparticles (SiNPs) and hybrid silica



**Fig. 1.** Schematic of different nanomaterials and biosensors used in detection infectious disease.

nanoparticles with solid or porous structures can be prepared via diverse methods such as hydrothermal, sol-gel, and reverse emulsion [70]. Silica nanoparticles can be further modified with organic dye which leads to rich color particles [71]. SiNPs are characterized by high stability, biocompatibility, chemical inertness, large surface area, transparency, and easy functionalization using silane chemistry [72,73]. Besides, silica can be used as a protective and stabilizing layer of magnetic nanoparticles or noble metal particles. Silica covering of gold particles is usually carried out with the Stöber method allowed thickness control with adjusting reaction time and reagent concentration [73]. A schematic of nanoparticles which are described in this section and the applied biosensors is illustrated in Fig. 1.

### 3. Biosensors for infectious disease based on nanomaterials

#### 3.1. Electrochemical biosensors

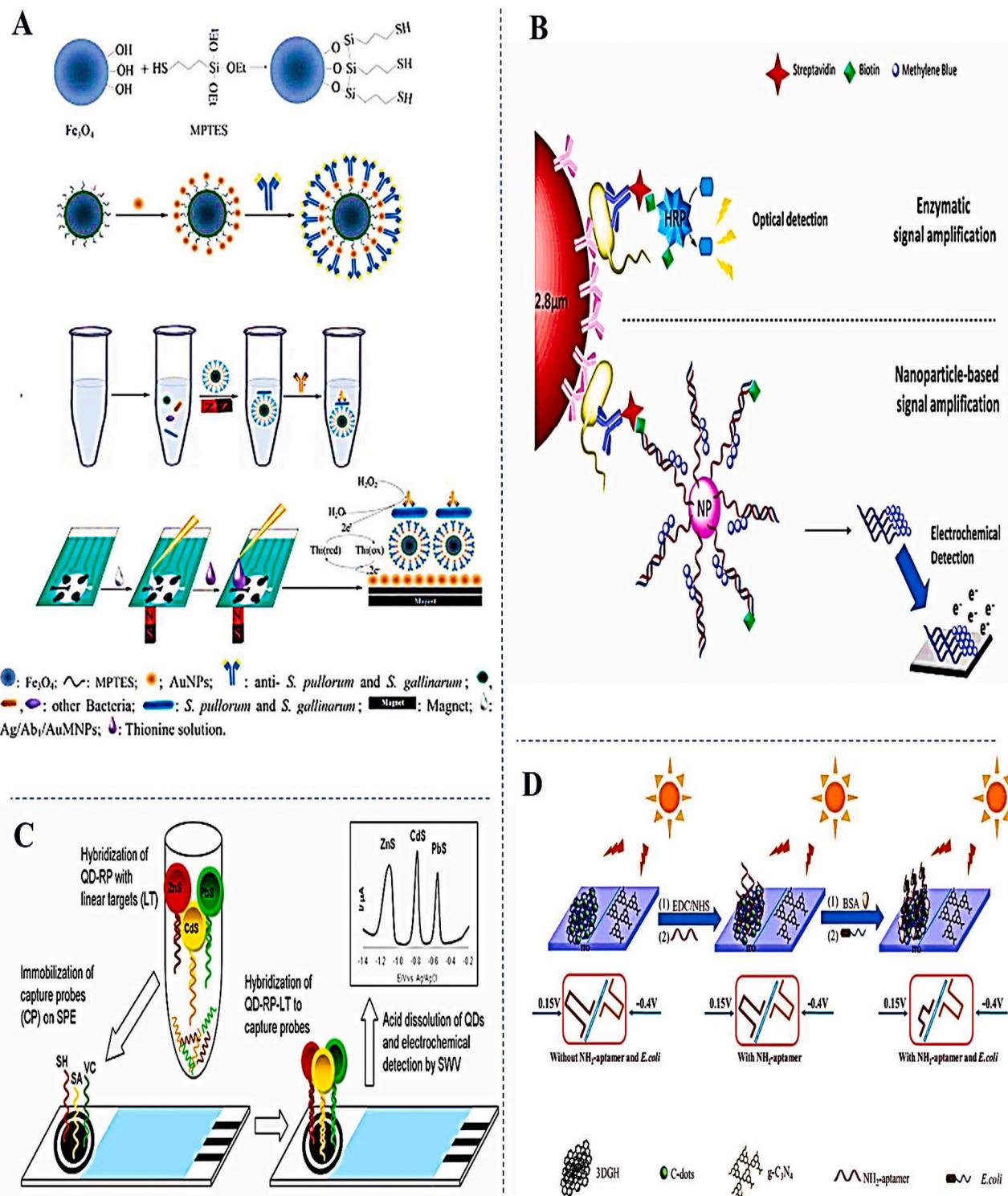
Electrochemical detection methods are widely used because of their easy procedure, high sensitivity, and ready to scale down for portable applications [74,75]. Also, screen printed technology facilitates the large scale fabrication of portable, low-cost analytical systems that can operate with small sample volume [76]. Nanomaterials have found extensive application in electrochemical sensors ranging from analyte separation/concentration (e.g. magnetic particle [77–90]) to the catalyst of redox reactions [23,90–102], enhancer of electrodes' surface conductivity [24,103], to grafting elements for efficient loading of biomolecules [79,93,104], as well as excellent sensitivity that achieve

with QDs [105–108].

Magnetic beads are commonly used in rapid assays to capture the targets from crude samples before detection [15]. The use of these not only allows to pre-concentrate the analyte (e.g. biomarkers) but also to reduce interferences that lead to an increase in the sensitivity and specificity of the sensors. Importantly superparamagnetic properties of magnetic nanoparticles allow their re-dispersion, with limited/no agglomeration, in the absence of the magnetic field. The surface of MNPs can be easily engineered, using specific coating providing in this way an appropriate functional group for the immobilization of biomolecules; furthermore, it has been shown that immobilization of biomolecules reduces their biodegradation when in contact with complex environmental or biological systems [78,90].

Electrochemical immunoassays have been effectively used for the detection of pathogens [78,90,91] due to simple handling methods, high sensitivity, and good applicability in real samples. Moreover, the sandwich assay enables the construction of a more specific and sensitive immunosensor in comparison to direct detection [23].

Fei et al. reported on the preparation of  $\text{Fe}_3\text{O}_4/\text{SiO}_2/\text{AuNPs}$  and their application in electrochemical immunoassaying.  $\text{Fe}_3\text{O}_4$ -NPs were prepared using the co-precipitation method; these, following capping with  $\text{SiO}_2$  and pre-functionalization with 3-Mercapto-propyltriethoxysilane (MPTES), were coated by AuNPs that served as anchoring element for *S. pullorum* and *S. gallinarum* antibodies. *S. pullorum* and *S. gallinarum* in the sample were captured by the  $\text{Fe}_3\text{O}_4/\text{SiO}_2/\text{AuNPs}$  and separated from the samples by applying an external magnetic field. Au-NPs/ $\text{Fe}_3\text{O}_4$ NPs with carrying bacteria, re-dispersed in buffer solution contained



**Fig. 2.** A) Schematic diagram of the synthesis process of the  $\text{Fe}_3\text{O}_4/\text{SiO}_2$  nanoparticle, immobilization antibody, capturing *S. pullorum* and *S. gallinarum* from sample, sandwich complex dropped on AuNPs/4-SPCE electrode with permission ref [90], B) Presentation of the assay with enzymatic amplification colorimetric detection and nanoparticle-based amplification and electrochemical detection with permission ref [15] C) Biosensor with npcRNA for simultaneous detection of three pathogens with permission of ref [105] D) Ratiometric photoelectrochemical aptasensor with permission [119].

horseradish peroxidase (HRP) labeled anti-*S. pullorum* and *S. gallinarum* to form a sandwich complex. The sandwich complex was dropped on 4-channel screen-printed carbon electrode already modified with electrodeposited AuNPs and supplied with a magnet, followed by the addition of thionine and hydrogen peroxide. To detect *S. pullorum* and *S. gallinarum* the reduction peak current change of the CV before and

after reaction of HRP with hydrogen peroxide was recorded. The antibody immobilized efficiently on the  $\text{Fe}_3\text{O}_4/\text{SiO}_2/\text{AuNPs}$  surface and 93.95% of the electrode signal was maintained after 30 days of storage indicated the ability of  $\text{Fe}_3\text{O}_4/\text{SiO}_2/\text{AuNPs}$  to retain the bioactivity of the adsorbed antibody [90]. The schematic of this procedure is depicted in Fig. 2A. Wang et al. [80] developed a portable electrochemical system,

based on a personal glucose meter (PGM), for detecting *E. Coli O157: H7* and *S. aureus*. In the reported work glucoamylase-quaternized magnetic nanoparticles (QMNP) were used to capture and detect bacteria, via a competitive mechanism. Following the capture of bacteria, via electrostatic interaction, on the positively charged QMNPs nanoparticles, Glucoamylase was released to the solution; this was then used to catalyze the hydrolysis of amylose, added in the solution into glucose which was then detected by PGM. QMNPs also prevented the growth of the bacteria which led to killing and monitoring pathogens at the same time [80]. Luo et al. used PGM to develop a rapid, portable, and sensitive immunosensor for *S. pullorum* and *S. gallinarum*. In this work capture of pathogens, performed using MNP-antibody, was followed by the introduction of an enzymatic label (antibody- SiNps-glucose oxidase (GOx)). The so generated sandwich system, following magnetic capture and cleaning, was dispersed in a glucose solution. Glucose concentration monitoring before and after hydrolysis of glucose by GOx revealed that the reduction in glucose concentration was proportional to the logarithm concentration of bacteria. The immune nanoparticles used in this work were shown to retain their activity after storage at 4 °C for at least 90 days [109].

Oxidoreductase Enzymes, like Horseradish peroxidase (HRP), are generally used in immunoassays to improve signal and reduce noise. Nanoparticles with peroxidase-like activity (e.g. Au@Pt bimetallic nanoparticle) are an attractive alternative to enzymes in immune assays resulting in more stable and cost-effective sensors [79,92]. For example, Zhu and coworkers fabricated two sandwich-type immunosensor with  $\text{Fe}_3\text{O}_4@\text{SiO}_2\text{-Ab1}/\text{E.coliO157: H7/rGO-NR-Au@Pt-Ab2s}$  for *Escherichia Coli O157: H7* detection [78,79]. One of the immunosensor utilized HRP [78] while the second one used Au@Pt nanoparticles instead of the enzyme [79]. The non-enzymatic system could reach a LOD of  $4.5 \times 10^2$  CFU mL<sup>-1</sup> and a linear range of  $4 \times 10^3$  to  $4 \times 10^8$  CFU mL<sup>-1</sup>; while the enzyme-based system was linear in  $4 \times 10^2$  to  $4 \times 10^8$  CFU mL<sup>-1</sup> concentration range with a LOD of 91 CFU mL<sup>-1</sup>. Besides, 91.0% of the initial response was retained after 5-weeks' storage of the Non-enzymatic electrode. Mo et al. also demonstrated the ability of non-enzymatic labels, reduced graphene oxide -natural red -Au@Pt, to retain their activity (85.46% of the initial signal) even after ten weeks of storage [92].

Gold nanomaterials have been widely used in electrochemical sensors due to biological compatibility, electrical conductivity, the high surface-to-volume ratio [22]. Moreover, gold nanomaterials can effectively couple with other nano/materials to produce various Au-based nanocomposites biosensor like poly(diallyldimethylammonium chloride)-functionalized graphene oxide and AuNps [98] chitosan/MWCNT/Polypyrrole/AuNPs [24] rGO/AuNR/poly thionine [110] to develop new electrochemical sensors with improving sensitivity, selectivity, and stability.

Xiang et al. developed a sandwich electrochemical immunosensor for the detection of *Salmonella* based on AuNPs dispersed in oxidized chitosan film on a glassy carbon electrode surface. In the reported work the presence of AuNPs in the film provided a more conducive platform for the immobilization of antibody, improved the catalytic activity of film for  $\text{H}_2\text{O}_2$  reduction and the immobilized HRP-antibody structure preserve high catalytic activity on the biocompatible substrate. The developed sensor presented good selectivity and reproducibility with higher sensitivity when compared to the sensor based on pure chitosan film [111]. Lin et al. compared two impedimetric biosensors, based on self-assembled thiolated protein G, which was used for oriented immobilization of goat anti - *E. coliO157: H7 IgG* using a bare gold electrode and gold nanoparticles modified electrode. Thiolated protein G was applied on the gold electrode surface while for the AuNP modified electrode, the gold electrode surface was first modified with a layer of 1, 6-Hexanedithiol, then colloidal AuNPs and thiolated protein G were applied on the surface. The AuNPs modified electrode was shown to have 2.2 times larger active area and a LOD 48 CFUmL<sup>-1</sup> for detection *E. Coli O157: H7* which is 3 times lower than those recorded for the bare

gold electrode [94]. Lin et al. reported on a regenerable and sensitive impedimetric immunosensor for the detection of type 5 adenovirus; the sensor was based on a multi-layered architecture assembled onto the surface of an Au electrode. The Au electrode was coated with AuNPs, grafted onto the sensor surface via 1, 6 hexane dithiol layers. 11-mercaptopundecanoic acid was then self-assembled on the sensor surface to attach monoclonal anti-adenovirus-5 antibodies. Regenerating the surface was carried out by applying a constant negative potential that could clean the gold electrode surface completely and enable the reuse of the electrode [95]. Ariffin and co-workers reported on ultrasensitive *E. coli* DNA biosensor based on a screen-printed carbon paste electrode modified with colloidal AuNPs and aminated hollow silica spheres (HSiSs) respectively. HSiSs was non-conductive and AuNPs promote the electron transfer from the intercalated DNA hybridization redox label to the SPE surface. HSiSs was treated with glutaraldehyde before attachment of aminated DNA. The sensor was able to detect target DNA in the  $1.0 \times 10^{-12} \mu\text{M}$  –  $1.0 \times 10^{-2} \mu\text{M}$  concentration range, with a low LOD of  $8.17 \times 10^{-14} \mu\text{M}$  via the DPV method; these performances were ascribed to the high loading of DNA probes taking place on the inner and outer surface of hollow silica, and to the conductivity increasing effect of AuNPs [96]. Bonnet and co-workers reported on a nanoparticle sandwich immunoassay based on the boron-doped diamond electrode for *E. coli* detection in platelet lysate. The system was based on solid-phase oligodeoxyribonucleotide (ODN) synthesis on nano-sized silica particles which was immobilized on micrometric silica beads. The nano-micro structures retained on the DNA synthesis column and ODN could synthesize on the silica nanoparticles with OH group via phosphoramidite chemistry. Then ODN-functionalized nanoparticles hybridized with methylene blue DNA probes. The methylene blue -ODN-functionalized nanoparticles were released from micrometric silica beads and used in the detection procedure. Capturing bacteria from the platelet lysate with antibody functionalized magnetic beads, followed by the formation of a sandwich complex by the second antibody with the streptavidin group. Then above mentioned methylene blue-ODN-functionalized silica nanoparticles, carried out the biotin group, coupled with sandwich structure via biotin/streptavidin chemistry. In the last step, the magnetic beads were suspended in RNase free water and the methylene blues labeled oligonucleotides were released. The boron-doped diamond electrode surface was grafted with a 4-aminobenzyl amine to produce a cationic surface. Then released methylene blue-labeled oligonucleotides were deposited on the surface while negative potential applied for electrostatic attachment. The square wave method was applied to detect the different concentrations of *E.coli*. The proposed system could achieve detection of  $10\text{--}100 \text{ CFU mL}^{-1}$  *E.coli* in platelet concentrates; this was 3 orders of magnitude more sensitive than the assay without nanoparticles. Moreover, the electrochemical system could provide 10 times superior LOD than that ELISA method with the same magnetic sandwich complex ELISA method which used HRP as a biocatalyst for TMB reduction [15]. The schematic of this assay is presented in Fig. 2B. DNA sensor for the invA gene of *Salmonella* was developed based on the combination of AgNCs (as a label) and cysteine-rich DNA reporting probes. Hybridization of target DNA and reporting DNA probes with the capture probe immobilized on AuNPs modified GCE electrodes resulted, following growth of the AgNCs (at the cystine groups of the reporting probes), in a clear DPV signal (oxidation of Ag). The sensor was demonstrated to have a linear response to DNA target concentration in a wide range between 1 fM and 0.1 nM with a LOD 0.162 fM [97]. Fei et al. reported on an immunoassay for the detection of *S. pullorum* and based on the combination of magnetic concentration and NP signal generation; more specifically AuNPs were used as redox labels in replacement of more conventional enzymatic labels.  $\text{Fe}_3\text{O}_4/\text{SiO}_2/\text{antibody1}$  captured *S. pullorum* from sample solution. After magnetic separation, the formed complex was incubated with/Ab<sub>2</sub>/AuNPs/rGO to produce a sandwich structure. The sandwich structure was dropped on the 4-SPCE electrode surface while applying magnet below the electrode. To generate an electrochemical signal

AuNPs were per-oxidation, by applying a constant potential followed by a DPV scan to reduce the in-situ generated  $\text{AuCl}^{-4}$  to  $\text{Au}^0$ . The recorded DPV signal correlated with the log CFU concentration of the bacteria and the system showed a LOD of  $89\text{CFU mL}^{-1}$  [77]. Nanocomposite of poly (diallyldimethylammonium chloride) graphene oxide (GO-PDDA) and AuNPs was reported in the manufacturing of an immunosensor for the detection of *E.coli* DH5  $\alpha$  in the dairy product. The nanocomposite was shown to provide a superior microenvironment for the immobilization of the antibody and 51.7% and 82.9% signal enhancement was achieved in comparison to sensor containing only GO or AuNPs. The sensor was shown to present a LOD 35 CFU  $\text{mL}^{-1}$  [98]. Valipour and Roushani investigated the application of AgNPs and thiol graphene quantum dot (AgNPs/GQD-SH) in the development of an immunosensor for the detection of hepatitis C virus core antigen. In the proposed sensor, the glassy carbon surface was first modified with the GQD-SH solution, and then AgNPs were attached via thiol groups of the GQD-SH/GCE surface. The antibody was immobilized on AgNPs by chemisorption between AgNPs and an amino group of anti-HCV core antigen. The electrochemical response was generated by the DPV monitoring of a redox probe (riboflavin). DPV currents decreased with the increase of the concentration of the antigen; The system was shown to allow to detect  $3 \times 10^{-12} \text{ g L}^{-1}$  of the hepatitis C virus core antigen with a broad linear range of  $5 \times 10^{-11}\text{--}6 \times 10^{-5} \text{ g L}^{-1}$  [112].

Pandey et al. proposed the use of a gold electrode decorated by reduced graphene oxide wrapped copper oxide and cysteine, in the development of *E. coli* O157: H7 immunosensor. Cysteine was used for the attachment of nanocomposite to the gold surface through the SH group and also as a green route for synthesis rGO. Antibody covalently bond via N-(3-dimethylaminopropyl)-N'-ethyl carbodiimide hydrochloride (EDC)/N-hydroxysuccinimide (NHS) coupling reagent through the carboxylic acid functional group of rGO. The presence of the Cu in the composite was shown to improve the LOD and sensitivity (slope of the calibration curve) of the sensor due to inducing stronger interactions between GO and Cysteine [113].

Bhardwaj et al. fabricated a paper-based electrochemical immuno-sensor that used carbon paste as a working electrode for the detection of *S. aureus*. EDC/NHS chemistry was utilized to activate carboxylic acid groups on the SWCNT surface and immobilization of the antibody, then SWCNT-antibody solution-dropped on the carbon paste surface. The DPV response of the sensor was shown to increase with the increase of the concentration of the bacteria; accordingly, to the authors, these results could be associated with three mechanisms: a) deficiency in antibody monolayer coverage as a result of bacteria binding, b) extracellular electron transfer of bacteria with redox mediator in solution and Ab-SWCNT, c) the presence of sodium ion on the bacteria cell wall [114]. Carboxylate graphene nanoflakes, prepared from MWCNT with wet chemical methods, were electrophoretically deposited on ITO coated glass substrate, and the so prepared surface used for the manufacturing a DNA sensor for the detection of a nucleic acid of *Escherichia coli* O157: H7. The developed genosensor presented higher sensitivity, ca. 2 order of magnitude when compared with resembling rGO counterpart sensor [115]. A dual aptamer sandwich complex consisting of a primary biotinylated aptamer (immobilized onto streptavidin magnetic bead) and a secondary aptamer conjugated with AgNPs was reported for the detection of *S. aureus*. The formed MB-Apt1/*S. aureus*/Apt2-AgNPs sandwich was collected and transferred in  $\text{HNO}_3$  solution (0.1 M) where AgNPs were dissolved to  $\text{Ag}^+$ . The reduction of the formed  $\text{Ag}^+$  was used as the analytical response of the sensor. Ag reduction signal was shown to be linear concerning *S. aureus* log concentration ( $10\text{--}10^6 \text{ CFU mL}^{-1}$ ); the sensor was reported to present a LOD of 1 CFU  $\text{mL}^{-1}$  [116].

A Nanoporous gold structure was manufactured on a glassy carbon electrode by the combination of electrodeposition of Au-Cu alloy and selective dealloying of Cu from the previous deposited film. The so prepared nanostructured surface was used, following attachment of thiolated *S. Typhimurium* aptamer on the surface, for the development of an impedimetric bacteria biosensor. The proposed sensor was shown to

present significant improvement, when compared to the planer Au electrode, in aptamer loading and surface stability. The aforementioned aptasensor was shown to present a low detection limit and excellent selectivity toward other bacteria and was demonstrated in the detection of *S. Typhimurium* in egg and rotten egg samples [99]. QDs embedded metal-organic framework (ZIF-8) particle were designed as signal-amplifying tags for the ultrasensitive and specific detection of *E. Coli* O157: H7. (CdS/ZIF-8) particles were modified with polyethyleneimine to introduce amino groups on their surfaces, before reaction with glutaraldehyde for immobilization *E. coli* O157: H7 antibody (CdS@ZIF-8/PEI/Ab). Glassy carbon electrode which was modified with p-aminobenzoic acid utilized to immobilize antibodies via EDC/NHS chemistry. The prepared electrode was incubated with bacteria and CdS@ZIF-8/PEI/Ab to form a sandwich complex on the electrode surface. Electrochemical detection of pathogens was made possible by the releasing of the Cd (II) ions from the CdS/ZIF-8 in HCl solution followed by their quantification via DPV. A LOD of  $3 \text{ CFU mL}^{-1}$  and a wide linear range  $10\text{--}10^8 \text{ CFU mL}^{-1}$  were reported; the sensor was also shown to be ca. 16 times more sensitive than the sensor using the CdS QD due to the large number of QD encapsulated in the MOF structure [117].

Vijian et al. reported on the multiplex detection of *Vibrio cholera*, *Salmonella* sp., and *Shigella* sp; in the reported work QDs (PbS, CdS, or ZnS) modified specific reporting probes were used to provide specificity to the sensor (QD-RP). Two hybridization strategies, the step-by-step and premix sandwich hybridization methods were applied for single and multiple pathogen detections. Premix sandwich hybridization methods had better results. For this strategy denaturing the PCR product of Non-protein coding RNAs (npcRNA) with different concentrations of three pathogens, were incubated with QD-RP conjugates. Then, Non-protein coding RNAs (npcRNA) QD-RP, one for each targeted DNA, were applied on SPE modified with a capture probe, followed by, QDs dissolution (using  $\text{HNO}_3$ ) and voltammetric detection, via SWV, of released  $\text{Zn}^{2+}$ ,  $\text{Cd}^{2+}$  and  $\text{Pb}^{2+}$  ions. Voltammetric detection of the heavy metal ions was made possible by the in-situ formation of a bismuth-film electrode. The prepared sensor presented the excellent LOD values of 22 aM (*Salmonella* sp.), 34aM (*Salmonella* sp.) and 42aM (*Shigella* sp) for single pathogen detection, and 51aM (*Salmonella* sp.), 53aM (*Salmonella* sp.), and 38aM (*Shigella* sp) for multiplex pathogen respectively [105]. The schematic of this procedure is illustrated in Fig. 2C.

Electrochemiluminescence aptasensor for the detection of *E.coli* using AgBr nanoparticles/3D nitrogen-doped GO hydrogel (AgBr/3DNGH) nanocomposite as the catalyst to enhance the electrochemiluminescence (ECL) of luminol was reported by Hao et al. Luminol was mixed with the AgBr/3DNGH before apply on the glassy carbon surface. The glassy carbon surface is then modified with a chitosan layer and aminated aptamer respectively. After capturing *E.coli* to the surface, ECL intensity was shown to decrease linearly in the  $0.5\text{--}500 \text{ CFU mL}^{-1}$  dynamic ranges; the sensor showed a LOD of  $0.17 \text{ CFU mL}^{-1}$ . The introduction of nitrogen in the GO structure was shown to improve the catalytic ability of the AgBr/3DGH composite; furthermore, the combination of AgBr and 3DNGH presented significantly improved catalytic performances when compared to the single material system [118].

A glassy carbon electrode was modified with electrodeposited Graphene oxide and decorated with CdTeQDs<sub>551</sub>-capture DNA<sub>HBV</sub> and CdTeQDs<sub>607</sub>-capture DNA<sub>HCV</sub> was used for the detection of HBV and HCV virus, followed by hybridization with the target probe. In the proposed biosensor AuNP-modified reporting probes, specific for the viruses were used as signal generation element. Virus detection was based on the ability of the AuNPs present in the specific reporting probes to quench the electrochemiluminescence of the CdTe QDs in an unreacted capture probe. The unreacted capture probe decreased with increasing concentration of target probe so the ECL signal was raised with increasing concentration of target probe in the range  $0.0005\text{--}0.5 \text{ nmol L}^{-1}$  and  $0.001\text{--}1.0 \text{ nmol L}^{-1}$  with LOD of 0.082 and  $0.34 \text{ pmol L}^{-1}$  for Hepatitis B and Hepatitis C virus respectively [106]. Hua and co-workers developed a sensitive potentiometric resolved ratiometric

**Table 1**

Electrochemical biosensors List of Abbreviation, Amperometry (AM), Antibody (Ab), (Aptamer (Ap), Au Nanoparticle citrate synthesis(AuNPsC), Biomolecule(Bio), Cyclic voltammetry (CV), Differential pulse voltammetry (DPV), Differential pulse anodic stripping voltammetry (DPASV), Electrochemiluminesense (ECL), Electrochemical Impedance Spectroscopy (EIS), Electropolymerized Au nanoparticle(AuNPsE), Horseradish peroxidase (HRP), Linear sweep voltammetry (LSV), Potentiometry (PT), Personal glucose meter(PGM), poly(diallyldimethylammonium chloride (PDDA), Square wave voltammetry (SWV), Square wave voltammetry anodic stripping voltammetry (SWVAW) [189–247].

Nanomaterial	Nanomaterial remark	Bi o	Pathogen	Met hod	Linear range	LOD	Remark	Real sample	Ref
AuNPs <sub>E</sub>	AuNPs 25nm, Ab attachment & enhancement of current of redox peak. Electrodeposited AuNPs coated electrode surface more uniformly & strongly	A b	<i>S. pullorum</i> & <i>S. Gallinarum</i>	CV	10 <sup>1</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	3.0 × 10 <sup>3</sup> CFU mL <sup>-1</sup>	Selective, Stable, Activity of Ab enhanced by Ionic Liquids as electrode modifier.	Egg Chicken Meat	[23]
AuNPs <sub>E</sub> , AuNPs <sub>C</sub> , Fe <sub>3</sub> O <sub>4</sub> /SiO <sub>2</sub> /AuNPs	AuNPs 25nm Enhancement electrochemical signal & attachment to Fe <sub>3</sub> O <sub>4</sub> /SiO <sub>2</sub> – SH Np with SH group Fe <sub>3</sub> O <sub>4</sub> MN co-precipitation method Fe <sub>3</sub> O <sub>4</sub> /SiO <sub>2</sub> /AuNPs 250 nm	A b	<i>S. pullorum</i> & <i>S. Gallinarum</i>	CV	10 <sup>2</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	3.2×10 <sup>3</sup> CFU mL <sup>-1</sup>	Selective, 4-SPE reduced the detection time & improves the reproducibility, Increase sensitivity by MNP	Chicken	[90]
GO-PDDA@AuNP	GO-PDDA@AuNP nanocomposites biocompatible& enhance electrochemical signal, AuNPs <sub>C</sub> 18nm, , Ab-Au-Thionine 108% increase in comparison with Ab-Thionine	A b	<i>E. Coli</i> (DH5α)	CV	50-5.0×10 <sup>6</sup> CFU mL <sup>-1</sup>	3.5 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Stable, Non- enzymatic, 90.8% of the response remained after 15 days	Fresh milk Yogurt Expired yogurt	[98]
Chitosan MWCNT Polypyrrole AuNPs	AuNPs increase the conductivity of the film	A b	<i>E. Coli</i> O157: H7	CV	3 × 10 <sup>1</sup> -3 × 10 <sup>7</sup> CFU mL <sup>-1</sup>	3×10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific, Disposable, 89.3% of signal remained after 2 weeks	-	[24]
rGO-NR-Au@Pt Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub>	Highest Reduction peak current when Pt/Au molar ratio is 1:1	A b	<i>E.coli</i> O157: H7	CV	4.0×10 <sup>2</sup> - 4.0×10 <sup>8</sup> CFU mL <sup>-1</sup>	91CFU mL <sup>-1</sup>	Specific, Non- enzymatic 91.0% response remained after 5 weeks.	Pork Milk	[78]
rGO-NR-Au@Pt Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub>	Au@Pt 30 nm, Fe <sub>3</sub> O <sub>4</sub> @ Solvothermal method 200 nm, Silica shell 30 nm Stöber method. Magnetic separation achieved in 10 seconds, rGO increase redox current	A b	<i>E.coli</i> O157:H7	CV	4 × 10 <sup>3</sup> -4×10 <sup>8</sup> CFU mL <sup>-1</sup>	4.5×10 <sup>3</sup> CFU mL <sup>-1</sup>	Specific, Au@Pt good peroxidase-like catalytic activity, Non- enzymatic 91.0% response remained after 5 weeks.	Pork Milk	[79]
rGO-NR-Au@Pt PANI AuNPs <sub>C</sub>	AuNPs work as a linker between PANI & Ab The AuNPs increase signal	A b	<i>E.coli</i> O157: H7	CV	8.9 × 10 <sup>3</sup> - 8.9 × 10 <sup>9</sup> CFU mL <sup>-1</sup>	2.84×10 <sup>3</sup> CFU mL <sup>-1</sup>	Specific, Non-enzymatic, Regenerable up to 5 times, 85.46 % responses remained after 10 week.	Milk & pork	[92]
Functionalized - MWCNT-Chit@Th	Thionine dye (Th) immobilized on functionalized-multi-walled carbon nanotube chitosan composite	A b	<i>Uropathoge</i> nic <i>E. coli</i>	CV	10 <sup>2</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	50 CFU mL <sup>-1</sup>	Selective, Easy preparation	Urine	[189]
Silica Np	Silica nanoparticle make quantitative measurement applicable	A b	<i>E.coli</i>	CV	10 <sup>2</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Fast, Easy	-	[190]
rGO–AuNPs –Ionic liquid	Electrochemical synthesis of rGO & AuNPs simultaneously. The composite showed better current response. AuNPs 100nm	A b	<i>E. Sakazakii</i>	CV	10 <sup>1</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	1.19×10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, RSD 6.2% n=7 3.9% signal change after 4 weeks.	Infant milk	[91]
AuNPs <sub>C</sub> -coated with reduced GO oxide, Fe <sub>3</sub> O <sub>4</sub> /SiO <sub>2</sub>	AuNPs & rGO simultaneously synthesised by citrate, AuNPs <sub>C</sub> 25nm capture Ab, rGO improve the peak current of the electrode, Fe <sub>3</sub> O <sub>4</sub> magnetic co-precipitation	A b	<i>S. Pullorum</i>	DP V	10 <sup>2</sup> - 0 <sup>6</sup> CFU mL <sup>-1</sup>	8.9×10 CFU mL <sup>-1</sup>	Selective, Non – enzymatic, AuNPs label signal	Chicken liver	[77]
GO/Au composite	AuNPs <sub>C</sub> 30 nm prepared in-situ in Go solution the intrinsic characteristics of GO preserved, Carboxyl groups in GO stabilizes the newly formed Au nanoparticles	A b	<i>C. sakazakii</i>	DP V	2.0 × 10 <sup>2</sup> -2.0 × 10 <sup>7</sup> CFU mL <sup>-1</sup>	2.0 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific, Synthesis of new antibody, Stable, 90% signal remained after 3weeks	Powdere d infant formula	[191]
Composite rGO/Thionine/ AuNPs	AuNPs <sub>E</sub> Composite formed with mixing rGO, Thionine & AuNPs. Thionine with positive charge linking between rGO & AuNPs	A b	<i>C. sakazakii</i>	DP V	8.8 × 10 <sup>1</sup> - 8.8 × 10 <sup>8</sup> CFU mL <sup>-1</sup>	1 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	Selective, 86.8% of the initial response remained after 30 days.	Infant milk powder Milk	[192]
Chitosan-AuNP composite film	AuNPs <sub>C</sub> , AuNPs/Chi has very good conductivity & demonstrates excellent electron-transfer ability. The optimum ratio of AuNPs/Chi is 1.	A b	<i>Salmonella</i>	DP V	10 <sup>-</sup> 10 <sup>5</sup> CFU mL <sup>-1</sup>	5 CFU mL <sup>-1</sup>	Selective, Simple fabrication	Tap water, Milk	[111]
AuNPs	AuNPs <sub>C</sub>	A b	<i>Salmonella</i>	DP V	100 - 700 Cell mL <sup>-1</sup>	100 cells mL <sup>-1</sup>	Fast, Low cost, disposable, Non-enzymatic	Skimmed milk	[193]
SWCNT	Peak current increased following the deposition of SWCNTs	A b	<i>S. aureus</i>	DP V	10 <sup>1</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	15, Milk 13 CFU mL <sup>-1</sup>	Low cost, RSD 10.5% after 5 weeks	Milk	[194]
PANI Coated MNP AuNPs	γ -Fe <sub>2</sub> O <sub>3</sub> solvothermal method nanoparticle core is 20 nm PANI-coated MNPs50 to 100 nm. AuNPs prepared in Alkaline solution with a dextrin capping agent.	A b	<i>E.coli</i> O157:H7	DP V	10 <sup>1</sup> - 10 <sup>6</sup> CFU mL <sup>-1</sup>	10 <sup>1</sup> CFU mL <sup>-1</sup>	Detection based on AuNPs signal	-	[81]
AuNR@SiO <sub>2</sub>	AuNRs with seed-mediated method AuNR@ SiO <sub>2</sub> . AuNRs 10 nm in mean diameter & about 40 nm in length	A b	<i>E.coli</i>	DP V	1.0×10 <sup>2</sup> -5.0×10 <sup>6</sup> CFU mL <sup>-1</sup>	60 CFU mL <sup>-1</sup>	Selective, Better LOD with AuNRs, 2% current response after 2 weeks.	Fresh milk Yogurt Expired yogurt	[195]
rGO/poly(etherelylene)mine	Electrodeposition rGO/PEI, rGO/PEI film formation cause a significant increase in redox current	A b	<i>E. coli</i> UTI89	DP V	10 <sup>1</sup> -10 <sup>4</sup> CFU mL <sup>-1</sup>	10 <sup>1</sup> CFU mL <sup>-1</sup>	Sensitive, 2.8 % loss peak current after 2 weeks storage	Urine Human serum	[196]
PPy-rGO/AuNPs	Deposited PPy on GO Surface prevent aggregation	A	<i>E. coli</i> K12.	DP	10 <sup>1</sup> - 10 <sup>7</sup>	1.0×10 <sup>1</sup>	Selective, 8.8% signal decrease after	Water	[197]

			b	V	CFU mL <sup>-1</sup>	CFU mL <sup>-1</sup>	40 days	Milk
PPy@Fc/AuNPs	of the GO& 6 times increase in surface area, AuNPs <sub>c</sub> 10 nm, GO & rGO decrease charge transfer resistance	A b	<i>E. Coli</i> <i>O157: H7</i>	DP V	7.8×10 <sup>1</sup> -7.8×10 <sup>6</sup> CFU mL <sup>-1</sup>	3.4×10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific, 92% of its initial response after 2 weeks	Spring water Milk [198]
Sulfonated rGO-PEDOT-AuNPs	AuNPs composites effectively facilitated the electron transfer, AuNPs 13 nm	A b	<i>E. coli</i> <i>O157: H7</i>	DP V	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	5.0×10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, 90.2% signal after 5 assay Milk Infant milk Powder Yogurt Expired yogurt	[199]
Poly(amidoamine) dendrimer-(PAMAM)encapsulated AuNPs/ CNT	AuNPs 3nm, HAuCl <sub>4</sub> : PAMAM ratio 55:1 the best ratio, Ab-CNT-HRP showed better DPV signal rather Ab- HRP, CNT -COOH 30nm	A b	<i>E. coli</i> <i>O157: H7</i>	DP V	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	300 cell mL <sup>-1</sup>	Specific	Beef [200]
AuNPs enclosed PANI	AuNPs 10nm prepared with mixing aqueous aniline & HAuCl <sub>4</sub> solution (353 K)	A b	<i>E. Coli</i> <i>O157: H7</i>	DP V	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	3CFU mL <sup>-1</sup>	Selective, ,16 times increase in CdS@ZIF-8 sensitivity rather than conventional sensor using CdS QDs,	Milk [117]
CdS quantum dots-encapsulated metal-organic frameworks as	DPV signals are amplified greatly by using CdS/ZIF <sub>8</sub> , CdS/ZIF <sub>8</sub> , with regular rhombic do-decahedron shape & an average diameter of 670 nm.	A b	<i>E. coli</i> <i>O157: H7</i>	DP V	10 to 10 <sup>8</sup> CFU mL <sup>-1</sup>	32CFU mL <sup>-1</sup>	Selective, Simultaneous detection	Stool water [201]
AuNPs,C60@AuN	AuNPs <sub>e</sub> , CdS 4nm PbS 10nm	A b	<i>E.coli</i> <i>O157: H7 &amp; V.cholerae O1</i>	DP V	5 × 10 <sup>1</sup> -1 × 10 <sup>6</sup> CFU mL <sup>-1</sup>	120 TCID <sub>50</sub> mL <sup>-1</sup>	Specific, 90.75% signal remained after 14 days	Normal avian serum [202]
Mesoporous graphitic-C <sub>3</sub> N <sub>4</sub>	mpg-C <sub>3</sub> N <sub>4</sub> has lower resistance than the bulk g-C <sub>3</sub> N <sub>4</sub> 12-nm used SiO <sub>2</sub> particle as a hard templet	A b	<i>Avian leukosis viruses J</i>	DP V	10 <sup>2.08</sup> 10 <sup>4.0</sup> TCID <sub>50</sub> mL <sup>-1</sup>	1.6 × 10 <sup>-9</sup> -1.6 × 10 <sup>-5</sup> g L <sup>-1</sup>	Selective, 91.2% signal retained after 30days	- [203]
AuNPs-rGO-Chitosan, AgNPs-rGO	Au <sup>3+</sup> & Ag <sup>+</sup> reduced to AuNPs by chitosan at 80°C.	A b	<i>Avian influenza virus H7</i>	LS V	1.6 × 10 <sup>-9</sup> -1.6 × 10 <sup>-5</sup> g L <sup>-1</sup>	6.8×10 <sup>9</sup> g L <sup>-1</sup>	Bi-MBs increase the detection signal 4 times.	Skim milk [204]
MNB	Superparamagnetic amino magnetic beads 200nm	A b	<i>Avian influenza A (H7N9) virus</i>	LS V	1 × 10 <sup>-8</sup> -2 × 10 <sup>-5</sup> g L <sup>-1</sup>	2.5×10 <sup>11</sup> -5 × 10 <sup>-1</sup> mole L <sup>-1</sup>	Dual sensor, Rapid response time, Simple fabrication	- [206]
GO-COOH MWCNT SiO <sub>2</sub> Np Chitosan	SiO <sub>2</sub> Np Sol-gel method, MWCNT significantly improve the electron transfer	A b	<i>HIV p24 Virus</i>	DP V	5. × 10 <sup>10</sup> -8.5 × 10 <sup>6</sup> g L <sup>-1</sup>	1.5×10 <sup>8</sup> g L <sup>-1</sup>	93.45%- 87.56% initial signal after 15 & 30 day	Human plasma [205]
Go	Go-methylene blue & chitosan	A b	<i>HA proteins of H5N1 &amp; H1N1</i>	DP V	2.5×10 <sup>11</sup> -5 × 10 <sup>-1</sup> mole L <sup>-1</sup>	5.2 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific, 95% separation efficiency due to MNP, Microelectrode reusable 50 time	Lettuce sample [207]
AgNPs GQD-SH	AgNPs <sub>c</sub> 13 nm, Graphene quantum dot-SH prepared from MWCNT & cystamine	A b	<i>Hepatitis C virus core</i>	DP V	5 × 10 <sup>-11</sup> -6 × 10 <sup>-5</sup> g L <sup>-1</sup>	3 × 10 <sup>12</sup> g L <sup>-1</sup>	90% signal remained after 20 days	Human serum [112]
			<i>antigen</i>					
Fe <sub>3</sub> O <sub>4</sub> COOH, AuNPs	MNP 180 nm AuNPs <sub>c</sub> 20nm	A b	<i>L.monocytogenes</i>	EIS	3.2 × 10 <sup>2</sup> -3.2 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	3.2 × 10 <sup>2</sup> CFU mL <sup>-1</sup>	Rapid, Easy to perform	Bovine milk [208]
Iron/gold core/shell & CdSe nanocrystals	Magnetic iron thermal decomposition method 10 ±3 nm & 15 ±3 nm due to the gold coating	A b	<i>Salmonella Typhimurium</i>	PT	10 <sup>2</sup> -10 <sup>7</sup> Cell mL <sup>-1</sup>	20 cells mL <sup>-1</sup>	Specific, Green synthesis AuNPs Ground beef 10 <sup>2</sup> -10 <sup>6</sup> CFU g <sup>-1</sup> LOD of 190 CFU g <sup>-1</sup>	Ground beef [82]
Fe <sub>3</sub> O <sub>4</sub> @poly(dopamine)	Fe <sub>3</sub> O <sub>4</sub> co-precipitation, Quasi-spherical with magnetic core 12nm, & 4nm polymeric layer,	A b	<i>L.pneumophila la</i>	AM	10 <sup>4</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	1 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	Specific, 30-day stability	Water [209]
Magnetic bead AuNPs	Magnetic bead 150 nm coated with GOx & dopamine (PMNCs), PMNCs capture 89% or bacteria cells in the range of 10 <sup>2</sup> - 10 <sup>5</sup> AuNPs glucose reduction method	A b	<i>E.coli O157: H7</i>	AM	10 <sup>2</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	5.2 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Electro catalytic properties AuNPs towards hydrogen	minced Beef Water [210]
AuNPs	AuNPs <sub>c</sub> superparamagnetic polystyrene beads	A b	<i>E. coli O157: H7 &amp; S. aureus</i>	AM	10 <sup>2</sup> -10 <sup>5</sup> CFU mL <sup>-1</sup>	148 CFU mL <sup>-1</sup>	Simple, Portable, Easy to use, Specific, without requiring expensive instruments. Immune nanoparticle stable for 90 days.	Milk [104]
AuNPs, AuNPs/HRP	AuNPs <sub>c</sub> 15 nm regular shape & uniform, AuNPs increase the fixed amount of HRP & the monoclonal Ab, & promote the electron transfer between the enzymes & the electrode surface	A b	<i>Salmonella</i>	AM	10 <sup>1</sup> -5 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	5 CFU mL <sup>-1</sup>	86.45% response signal after 13 day, Preparation & purification of new Salmonella plasmid virulence antibody	[109]
Magnetic nanoparticle	Magnetic nanoparticle illustrated an obvious higher matrix effect & higher agglomeration effect but better LOD After 8h enrichment	A b	<i>Salmonella</i>	AM	598 for micro particle & 291 for nanoparticle CFU mL <sup>-1</sup>	598 for micro particle & 291 for nanoparticle CFU mL <sup>-1</sup>	Simple, Portable, Easy to use, Specific, without requiring expensive instruments. Immune nanoparticle stable for 90 days.	Milk [83]
SilicaNp, MNP	Antibody- MNPs co-participation method, MNPs10nm, narrow size distribution, Magnetic separation obtained in 1 min, Silica nanoparticle 50 nm inverse microemulsion	A b	<i>S. Pullorum</i> <i>S.Gallinarum</i>	PG M	1.27 × 10 <sup>2</sup> -1.27 × 10 <sup>5</sup> CFU mL <sup>-1</sup>	7.2 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Very good stability Portable, Low cost, Easy to use Ab-Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> NPs & Ab/invertase-Au@Pt/SiO <sub>2</sub> NPs stable for 90 days.	[109]
Au@Pt/SiO <sub>2</sub> NPs Fe <sub>3</sub> O <sub>4</sub> NP/SiO <sub>2</sub>	poly-(4-styrene sulfonic acid- co -maleic acid) (PSSMA) as a linker between Au@Pt & SiO <sub>2</sub> MNP 300 nm spherical nanocrystal solvothermal method clusters, Silica thickness 50 nm	A b	<i>E. coli O157: H7</i>	PG M	3.5 × 10 <sup>2</sup> - 3.5 × 10 <sup>8</sup> CFU mL <sup>-1</sup>	1.83 × 10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, Portable, recoveries from 80% to 94.8% Portable	Milk [84]
ST-magnetic nanobeads	Streptavidin magnetic nanobeads size 150nm	A b	<i>E. coli O157: H7</i>	PG M	10 <sup>2</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	79 CFU mL	Milk [85]	

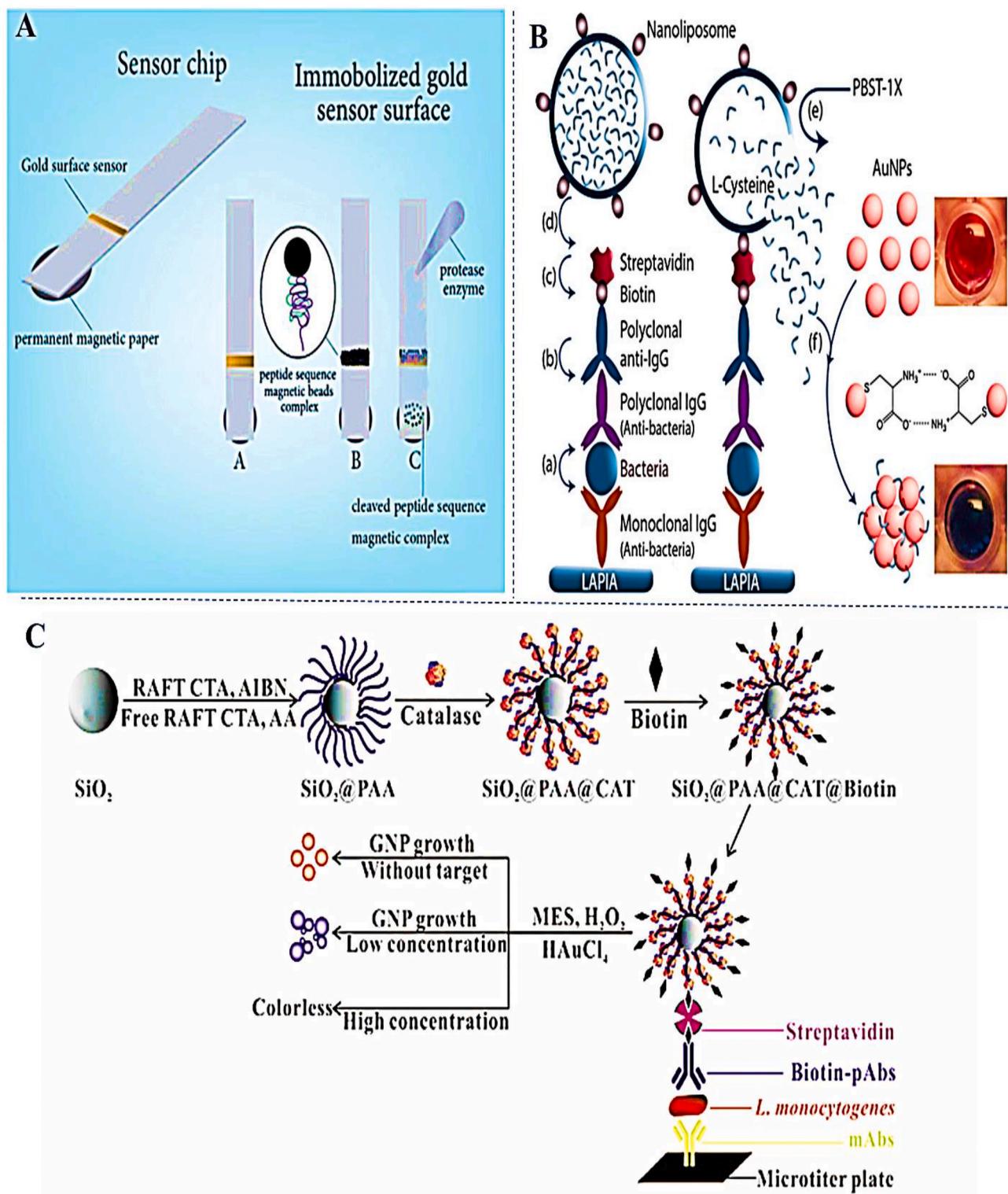
Fe <sub>3</sub> O <sub>4</sub>	Hydrothermal, MNPs 200 nm are regular spherical particles, Good separation in a few second	A b	<i>E. coli</i> <i>O157: H7</i>	PG M/I CA	2.08 × 10 <sup>2</sup> – 2.08 × 10 <sup>8</sup> CFU mL <sup>-1</sup>	1.04 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	Selective, Portable ,	Milk	[86]
Magnetic nanoparticle, Silica Np	Fe <sub>3</sub> O <sub>4</sub> Co-precipitation 10nm coated with silica layer 2nm,60 s separation, SilicaNp reverse microemulsion method uniform particle 50 nm	A b	<i>C.sakazakii</i>	PG M	9 × 10 <sup>2</sup> – 9 × 10 <sup>7</sup> CFU mL <sup>-1</sup>	4 × 10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Portable, 100-70% signal retained after 1- 90 days	Powdered infant milk	[87]
AuNPs	AuNPs 10nm, AuNPs enhance the transfer of electrons, The electroreactive surface area of the Au NP-modified electrode 2.2 times of the planar gold electrode	A b	Heat killed <i>E. coli</i> <i>O157: H7</i>	EIS	10 <sup>2</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	48 for AuNPs & 140 CFU mL <sup>-1</sup> planer electrode	Selective, AuNPs 3 time lower LOD	-	[94]
GO-wrapped copper oxide-cysteine	Cu-doped rGO Cysteine facilitated strong adsorption leading to faster electron transfer kinetics in comparison to that of the electrode without GO	A b	<i>E. coli</i> <i>O157: H7</i> <i>E. coli</i> <i>DH5α</i>	EIS	10 <sup>1</sup> –10 <sup>8</sup> CFU mL <sup>-1</sup>	3.8 CFU mL <sup>-1</sup>	Selective, 90.2% signal remained after 40 days, 8 times regeneration, Cu in composite structure increase the sensitivity & LOD	Tap Water, Juices, Skimmed milk	[113]
Streptavidin Fe <sub>3</sub> O <sub>4</sub>	Streptavidin Fe <sub>3</sub> O <sub>4</sub> 150 nm	A b	<i>E. coli</i> <i>O157: H7</i>	EIS	10 <sup>4</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	1400 cells	1400 cells in less than 1 hour.	Ground beef	[211]
G/GO	Composite of Graphene/Graphene oxide GO resulted in an enhanced conductance value.	A b	<i>S.Typhimurium</i>	EIS	10 <sup>1</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific, Using a specific antibody against specific surface antigen OmpD for the first time	Lichi & orange juices	[212]
AuNPs	AuNPs 10nm Au NPs promoted electron transfer between the electrode surface & the redox probe	A b	<i>Type5 Adenovirus</i>	EIS	10 <sup>1</sup> –10 <sup>8</sup> virus particle mL <sup>-1</sup>	30 virus particle mL <sup>-1</sup>	Selective, Sensitive, 30 times regeneration RSD 5.1%	-	[95]
SilicaNps –MB-DNA	Rhodamine encapsulated in the core of SilicaNps 50 nm	A b	<i>E. coli</i>	SW V	10 <sup>1</sup> –10 <sup>4</sup> CFU mL <sup>-1</sup>	1×10 <sup>1</sup> CFU mL <sup>-1</sup>	The Nps amplify sensitivity 1000 times compared with the assay run without NPs	Platelet concentrates	[15]
HA-NH2-CNT	Stable film	A b	<i>Hepatitis B virus core protein</i>	SW V	1·6×10 <sup>6</sup> g L <sup>-1</sup>	3×10 <sup>8</sup> g L <sup>-1</sup>	Easy synthesis	-	[213]
AuNPs MNP	PANI coated MNP 50-100 nm, AuNPs-conjugated PbS nanoparticles & capture antibody	A b	<i>E. Coli</i> <i>O157: H7</i>	SW AS V	10 <sup>1</sup> –10 <sup>6</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Optimize antibody to maintain colloid stability, 94.96% bacteria capture with MNP	-	[214]
Au Nanorod	AuNRs 32.8 ± 1.8 nm (length) & 6.7 ± 1.2 nm width, growth seed method, Rct sharply decreased after AuNRs deposition	T4-bacteriophages	<i>E. coli K.12</i>	EIS	-	10 <sup>3</sup> CFU mL <sup>-1</sup>	Low cost	-	[215]
polyethylenimine (PEI)-functionalized carbon nanotube	Functionalized CNT directly grown on the surface caused uniform surface deposition of oriented phage & immobilization with minimal particle aggregation.	T2-bacteriophage	<i>E.coli B</i>	EIS	10 <sup>3</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	1.5×10 <sup>3</sup> CFU mL <sup>-1</sup>	Specific, Induced immobilization methods result in 15 X higher loading of phage particles on the electrode surface	-	[216]
(CNTs)	Antimicrobial peptide clavanin A	<i>K.pneumoniae</i> , <i>E. faecalis</i> , <i>E. coli</i> , <i>B. subtilis</i>	-	EIS	-	1.0×10 <sup>2</sup> – 1.0×10 <sup>6</sup> CFU mL <sup>-1</sup>	Differentiate between gram-positive & gram-negative bacteria, High sensitivity to gram-negative bacteria	-	[217]
AuNPs	AuNPs <sub>c</sub> 15nm	-	DNA probe <i>Vibrio cholerae</i>	DP V	4 to 10 <sup>3</sup> CFU mL <sup>-1</sup>	2 CFU mL <sup>-1</sup>	Selective	Stool	[218]
AuNPs/hollow silica spheres	AuNPs- doped with aminated HSIs exhibited high electron transfer performance, AuNPs for increasing the biosensor conductivity	-	DNA probe <i>E. coli DNA</i>	DP V	1×10 <sup>18</sup> –1 × 10 <sup>8</sup> mol L <sup>-1</sup>	8.17 × 10 <sup>-20</sup> mol L <sup>-1</sup>	Regenerable 4 times	River water	[96]
AuNPs <sub>E</sub> , AgNCs+ AuNPsC	Peak currents increased after the electrodeposition of AuNPs <sub>E</sub> , AuNPs carried capture probe. AuNPs cause loading more ssDNA, AgNCs work as the signal inducer	-	DNA probe PCR product of <i>Salmonella</i>	DP V	1×10 <sup>15</sup> –1 × 10 <sup>11</sup> mol L <sup>-1</sup>	1.62 <sup>-16</sup> mol L <sup>-1</sup>	DPV responses maintained 88.2% & 71.4% of its initial signal response respectively after 5 & 10 days of storage, invA gene sequence of <i>Salmonella</i> in PCR samples	-	[97]
Au nanoaggregate	AuNAs prepared at physiological pH enhance the electrochemical signal &cause high loading of the ssDNA probe. Crystallite size of AuNAs 7.9 nm & 13.5.	-	Salmonella specific DNA str& <i>S. Typhi</i>	DP V	4×10 <sup>18</sup> –2.4×10 <sup>16</sup> mol L <sup>-1</sup>	Selective, stability 60 Day &reusability of 6 time, signal 8 % loss after 60 days, The electroactive area & greater for AuNAs-ITO than for bare ITO.	-	Urine & blood	[93]
rGO & NiO nanoparticle	Electro reduction of GO, Redox peak current increased after Go reduction on the surface, Electrodeposition of NiO nanoparticle, NiO 125 nm	-	Selected part <i>S.enteritidis</i> gene	DP V	1.0 × 10 <sup>-13</sup> -1.0 × 10 <sup>-6</sup> mol L <sup>-1</sup>	3.12×10 <sup>-14</sup> mol L <sup>-1</sup>	Selective, 96 .3 % of the initial after 10 day PCR product of gene sequence	-	[219]
PPy- rGo, AuNPs <sub>E</sub> , AuNPs-HRP-SA	AuNPs <sub>E</sub> , AuNPs <sub>C</sub> size 10 nm, Chemical reduction of GO,PPy- rGo & AuNPs <sub>E</sub> increased peak current. Signal decrease without using AuNPs	-	<i>S. Typhimurium</i> strains <i>invA</i> gene <i>S. Typhimurium</i>	DP V	1.0 × 10 <sup>-16</sup> -1.0 × 10 <sup>-10</sup> mol L <sup>-1</sup> 9.6–9.6×10 <sup>4</sup> CFU mL <sup>-1</sup>	4.7×10 <sup>-17</sup> mol L <sup>-1</sup> 8.07 CFU mL <sup>-1</sup>	Sensitive, Good regeneration ability 91.6 % signal after 3 cycle regeneration, 84.6% original signal after 7-day storage, PCR product of salmonella	-	[220]
MWCNT	More sensitive sensor with MWCNT	-	DNA probe <i>16S rDNA</i> for <i>E. coli</i>	DP V	7-12 × 10 <sup>3</sup> g L <sup>-1</sup>	1.7×10 <sup>-8</sup> mol L <sup>-1</sup>	Selective PCR amplified products	-	[221]
AuNPsE, AuNPsC	Flower-like CNTs-PAN nanohybrids possessed large surface area, excellent electronic conductivity, & redox activity, Fullerene nanoparticle prepared by mixing poly(amino-amine)dendrimers & fullerene.	-	IS6110 DNA the sequence of Mycobacterium tuberculosis M. tuberculosis	DP V	1×10 <sup>15</sup> –1×10 <sup>8</sup> mol L <sup>-1</sup>	3.3×10 <sup>-16</sup> mol L <sup>-1</sup>	Selective, Current response of five electrodes retained from 94.5% to 96.9% after 10 days storage	Sputume	[222]
rGO/AuNR/poly thionine	AuNRs with seed growth mediated method rGO/Au NR/PT excellent electric conducting material & accelerated the electron transfer.	-	DNA probe Human papillomavirus	DP V	1×10 <sup>13</sup> –1 × 10 <sup>5</sup> mol L <sup>-1</sup>	4.03×10 <sup>-14</sup> mol L <sup>-1</sup>	Selective, Sensitive, Wide linear range	Human serum	[110]

(HPV)							
Graphene-COOH nanoflakes, rGO	COOH Graphene nanoflakes prepared from MWCNT 50nm with a wet chemical method	DNA probe <i>nucleic acid Escherichia coli O157: H7</i>	EIS	$10^{-17}$ - $10^{-6}$ mol L <sup>-1</sup>	$1 \times 10^{-17}$ mol L <sup>-1</sup>	15% Loss after the 6th cycle of regeneration, The 18% decrease in the signal response after 30 days	[115]
GO-iron oxide-chitosan hybrid nanobiocomposite	Electrophoretic deposition GO-iron oxide-chitosan monodispersed spherical shape 8 to 12 fast electron transfer kinetics	DNA probe <i>E.coli</i>	EIS	$10^{-14}$ - $10^{-6}$ mol L <sup>-1</sup>	$1 \times 10^{-14}$ mol L <sup>-1</sup>	Selective, 10.37, 10 % loss of the initial value after 6 cycles of regeneration &after 6-week storage	[223]
AuNPs	AuNPs <sub>C</sub> 16 nm	DNA probe, <i>Leishmania DNA</i>	Am	0.5-500 parasites mL <sup>-1</sup> blood	0.8 parasites mL <sup>-1</sup> blood	AuNPs primer 8 week lifetime	Dog Blood Parasite [224]
GO-Tetracycline ZrO <sub>2</sub> Np	ZrO <sub>2</sub> increased the conductivity of the electrode	DNA sample, <i>Mycobacterium avium subspecies paratuberculosis DNA</i>	EIS	$6.37 \times 10^{15}$ M (2 $\times 10^{14}$ g/L) $6.37 \times 10^{15}$ M (20 $\times 10^8$ g/L)	$6.37 \times 10^{15}$ M (2 $\times 10^{14}$ g/L)	Rapid	Faecal sample Cattle [225]
SiNPs doped MB	Silica Np reverse microemulsion method, SiNPs/ MB 100 nm	Viral DNA, hepatitis C virus DNA	EIS	$100$ - $10^6$ Copies mL <sup>-1</sup>	90 Copies mL <sup>-1</sup>	Selective, Regenerable, 10-day stability	Serum [226]
QD	Quantum dot reverse micelles, the narrow absorbance peak of high mono dispersion of QDs in aqueous solution	non-protein-coding RNA, sequences ncRNA sequences of Vibrio cholera, Salmonella sp. & Shigella sp	SW AS V	$1-6 \times 10^{17}$	$2.2 \times 10^{17}$ (VC-PbS), $3.4 \times 10^{17}$ , (SA-CdS) & $4.2 \times 10^{17}$ (SH-ZnS) mol L <sup>-1</sup>	Simultaneously detection npcRNA sequences of Vibrio cholerae, Salmonella sp., & Shigella sp in the attomolar range.	PCR product [105]
Porous AuNPs	AuNPs strongly facilitate the electron transfer & provides the higher surface roughness & the active area which higher coverage, Ag nanoplates as a template for seed-growth method 60–110 nm diameter distribution	multi-functional DNA hemagglutinin (HA) protein Ap + HRP-DNAzyme with hemin HA protein in AIV H5N1	CV	$10^{12}$ - $10^7$ mol L <sup>-1</sup>	$10^{12}$ mol L <sup>-1</sup>	Selective	Chicken serum [101]
MoS <sub>2</sub> -rGO nanocomposite	rGO avoid restacking MoS <sub>2</sub> nanocomposite .	A p V <sub>i</sub> polysaccharide antigen mediated detection of enteric fever	SW V	$10^{10}$ - $10^6$ g mL <sup>-1</sup>	$10^{10}$ g mL <sup>-1</sup>	Selective, New aptamer	Blood, Urine [227]
AuNPs	AuNPs <sub>E</sub> Capture probe, AuNPs increase LOD & Linear range	A p S. <i>Typhimurium</i>	DP V	$20-2 \times 10^8$	16 CFU mL <sup>-1</sup>	Selective	Bottled mineral water [228]
AgNPs	AgNPs 20 nm prepared by NaBH <sub>4</sub>	A p S. <i>aureus</i>	DP AS V	$10^1$ - $10^6$ CFU mL <sup>-1</sup>	1 CFU mL <sup>-1</sup>	Excellent sensitivity, Specific	Tap & river water [116]
AuNPs <sub>C</sub> , AgNPs Hollow gold/silver nanospheres, Ag@Au core-shell, Au@Ag core-shell, Ag/Au alloy	Ag@Au better result core enhance the physical properties & the shell offers chemical stability & biocompatibility for the immobilization of aptamers, 30 to 100 nm	A p E. <i>coli</i>	DP V	$10^1$ - $10^5$ CFU mL <sup>-1</sup>	90 CFU mL <sup>-1</sup>	Comparison of different nanoparticle	[229]
AuNPs, Uio-67/GR	AuNPs <sub>C</sub>	A p S. <i>Typhimurium</i>	DP V	$2 \times 10^1$ - $2 \times 10^8$ CFU mL <sup>-1</sup>	5 CFU mL <sup>-1</sup>	Selective, 93.1% current reserved after 15 days n=6	Milk [230]
Nanocomposite of rGO -azophloxine	Chemical reduction of Go, GO-Ap formed by $\pi$ - $\pi$ stacking. Doping of rGO with AP dye improves its free charge carrier density	A p S. <i>Typhimurium</i>	DP V	$10^1$ - $10^8$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Fast 5 min incubation time, Low cost, 5.1 % increase in current response after 20day	- [54]
rGO/CNT	rGO-CNT Hydrothermal method, Hybridization of rGO with carbon nanotubes (CNT) avoid the aggregation of graphene sheets because of Working CNT as a spacer	A P S. <i>Typhimurium</i>	DP V	$10^1$ - $10^8$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Selective, 20-day stability,	Chicken [231]
rGO-TiO <sub>2</sub>	Chemical reduction of Go, Noticeable increase in current density observed after coating rGO-TiO <sub>2</sub> compared with Go.	A p <i>Salmonella enterica</i>	DP V	$10^1$ - $10^8$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Selective, Fast 5min incubation, Stability to 21 days, & a 10% decrease in signal after 30 days.	Chicken [232]
AuNPs <sub>E</sub> , AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 18 nm spherical & homogeneously distributed AuNPs on the electrode surface enhanced the peak current 2 times	A p <i>Salmonella</i>	DP V	$2 \times 10^1$ - $2 \times 10^6$ CFU mL <sup>-1</sup>	$20$ CFU mL <sup>-1</sup> $200$ CFU mL <sup>-1</sup> in milk	Selective, Easy to use	Milk [233]
rGo + chitosan	Electro reduction Go-CHI solution, Current increased after RGO-CHI deposition	A p <i>Salmonella</i>	DP V	$10^1$ - $10^6$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Selective, Thiol-aptamer gave a higher current compared to amine aptamer, Specific	Chicken [234]
SWCNT	SWCNT attachment of aptamer with $\pi$ - $\pi$ stacking	A p <i>Escherichia coli O157:H7</i>	DP V	$1.7 \times 10^1$ - $1.1 \times 10^7$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Selective	Tap water [235]
Go	Methylene blue anchored Go	A p <i>C. sakazakii</i>	DP V	$2 \times 10^1$ - $2 \times 10^6$ CFU mL <sup>-1</sup>	$7$ CFU mL <sup>-1</sup>	Specific, 88. 4% of its original response after 15-day storage,	Powdered infant milk [236]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> thiocyanuric acid/gold nanoparticles worked as the amplifier	A p <i>Avian influenza virus H5N1</i>	EIS	0.25 -16 HAU	0.25 HAU for the pure virus & 1 HAU for the tracheal	Specific, 57-fold increase for the 1 HAU samples with an amplifier	Swab samples of chicken [237]

Chicken swap										
Go/silica-cobalt	GO/Si-10Co has a small size while rGO/Si-10Co has mesopores with a large size. rGO/Si-10Co Showed a higher surface area.	A p	Salmonella	EIS	$10^1\text{-}10^8$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	5 min incubation time with bacteria		[238]	
Nanoporous gold	Nanoporous gold prepared by electrodeposition of Au-Cu alloy at the surface of the electrode & dealloying of Au-Cu film from Cu.	A p	Salmonella	EIS	$6.5\times10^2\text{-}6.5\times10^8$ CFU mL <sup>-1</sup>	1CFU mL <sup>-1</sup>	New, facile, & cost-effective nanoporous gold Synthesis, Regenerable, Distinguish between live & dead bacteria cells.	Egg, Rotten egg	[99]	
rGO & COOH-MWCNT	Electrodeposition GO-MWCNT & electrochemical reduction to rGO-MWCNT GO oxide with a uniform length	A p	Salmonella	EIS	75 to $7.5\times10^5$ CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup>	Selective, The current of rGO -MWCNT is bigger than MWCNT alone		[239]	
MWCNT -COOH	Electrodeposition MWCNT increased peak current times.	A p	S.Typhimurium & S.enteritidis	EIS	$10^1\text{-}10^5$ CFU mL <sup>-1</sup>	$6.7\times10^1\& 5.5\times10^1$ CFU mL <sup>-1</sup>	Selective, Sensitive, Low cost of preparation	Chicken	[240]	
GO -Platinum nano cauliflower hybrid	Chemical reduction of GO, Pt was electrodeposited on GO-nano cellulose, Pt nano cauliflower structures ranging from 10 nm nodes to 2 $\mu\text{m}$ .	A p	Escherichia coli O157: H7	EIS	4 to $10^3$ CFU mL <sup>-1</sup>	4 CFU mL <sup>-1</sup>	POC biosensing, Electroactive surface area (ESA) increased by an order of magnitude after deposition of Pt nano cauliflower structures		[241]	
MWCNT-COOH	Bridged Rebar Graphene synthesized from MWCNT using modified chemical assisted unscrolling method followed by bridging with terephthalaldehyde	A p	E. coli O78:K80: H11	EIS	$10^1\text{-}10^6$ CFU mL <sup>-1</sup>	$10^1$ CFU mL <sup>-1</sup>	Specific, New aptamer	Water Juice Milk	[242]	
AuNPs, Carbon nanoparticles/ cellulose nanofibers	Significant increase in the peak current & decrease in Rct CNFs with AuNPs/CNPs nanocomposite.	A p	S. aureus	EIS	$1.2\times10^1\text{-}1.2\times10^8$ CFU mL <sup>-1</sup>	1 CFU mL <sup>-1</sup>	Selective, Discriminate between live and dead bacteria cells	Human blood serum	[243]	
AgNPs	AgNPs <sub>100</sub> nm modification of GCE by AgNPs enhanced electron -transfer of modified electrode & reduce charge transfer resistance	A p	P. aeruginosa	EIS	$10^1\text{-}10^6$ CFU mL <sup>-1</sup>	33 CFU mL <sup>-1</sup>	Selective, First impedimetric for P. aeruginosa	Human blood serum	[102]	
GQD	Aptamer molecules attached on the surface of GQD by chemisorption between GQD & amine groups of aptamer by $\pi$ -stacking interactions	A p	hepatitis C virus ( HCV) core antigen	EIS	$10\text{-}70\times10^{-9}\text{ g L}^{-1}$ & $7\text{-}40\times10^{-8}\text{ g L}^{-1}$	$3.3\times10^{-9}\text{ g L}^{-1}$	96-85% initial signal after 10 &14 days	Human serum	[244]	
AgBrNPs anchored 3D nitrogen-doped GO hydrogel	AgBr/3D N-doped GO hydrogel, AgBr nanoparticles was around 50–180 nm, Smaller impedance of AgBr/3DNGH due to presence of rGO	A p	E.coli	EC L	0.5 to 500 CFU mL <sup>-1</sup>	0.17 CFU mL <sup>-1</sup>	Simple operation process, low-cost, 12 Day storage time	Meat	[118]	
AuNPs <sub>E</sub> , rGO	Chemical reduction of GO, AuNPs <sub>E</sub> rGO & AuNPs increase electrical conductivity & electron	A p	S. aureus	EC L	$1.5\times10^2\text{-}1.5\times10^8$ CFU mL <sup>-1</sup>	28 CFU mL <sup>-1</sup>	Selective, Enzyme free High ECL with AuNPs &	Urine	[245]	
MoS <sub>2</sub> -PtNPs	transferrability						catalytic activity of hemin			
GO COOH/porcine IgG	GO-COOH strong immobilization ability to porcine IgG for its large surface & the high number of carboxyl groups. COOH- GO facilitated electron transfer	S. aureus protein A S. aureus	EC L	$1.0\times10^3\text{-}1.0\times10^9$ CFU mL <sup>-1</sup>	$3.1\times10^2$ CFU mL <sup>-1</sup>	Selective, Simple fabrication, Short assay timer. 91.3% of initial signal remained for 15 days.	Milk, Human Saliva		[246]	
Ru(bpy) <sub>3</sub> <sup>2+</sup> -PtNps complex	Fe <sub>3</sub> O <sub>4</sub> @ Au NPs 50nm was more suitable, Ru-PtNps 12nm, rGO & Ru-PtNps decreased charge transfer resistance	A b Fe <sub>3</sub> O <sub>4</sub> @Au Nps, rGO	Clostridium perfringens	EC L	$10^1\text{-}10^7$ CFU mL <sup>-1</sup>	$10^7$ CFU mL <sup>-1</sup>	Specific, POC analysis	Infected Wound	[88]	
CdTe QDs, AuNPs, GO	AuNPs <sub>C</sub> , Electrochemical reduction of GO	DNA probe Hepatitis B virus& Hepatitis C virus DNA	EC L	$5\times10^{13}\text{-}5\times10^{10}$ $1\times10^{12}\text{-}1\times10^9$ mol L <sup>-1</sup>	$8.2\times10^{14}$ Hepatitis C virus $3.4\times10^{13}$ mol L <sup>-1</sup>	Sensitive, Multiplex sensing. 92% signal value remained after 15 days	Human Serum		[106]	
Fe <sub>3</sub> O <sub>4</sub> /Van/PEG	Amin terminated Fe <sub>3</sub> O <sub>4</sub> 100 – 200 nm Vancomycin coated	Bacteria DNA L. monocytogenes	EC L	$10^2\text{-}10^6$ CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective, Fe <sub>3</sub> O <sub>4</sub> /Van/PEG Biocompatible & 90% Capture bacteria			[89]	
Fe <sub>3</sub> O <sub>4</sub> @GO	Fe <sub>3</sub> O <sub>4</sub> Co-Precipitation method	A b Fe <sub>3</sub> O <sub>4</sub> @GO	V.parahaemolyticus	EC L	$10\text{-}10^8$ CFU mL <sup>-1</sup>	5 CFU mL <sup>-1</sup>	Specific, $92.97\pm4.5\%$ initial signal n=5 after 1 month, High & stable ECL signal, RSD 7.8% n=5, Recoveries 94.4–112.0%, Simple instrumentation(a laboratory built ECL)	Seafood, Seawater		[247]
Hydrogel-loaded carbon quantum dots(Cdots/3DGH) &Graphene-like carbon nitride (g-C <sub>3</sub> N <sub>4</sub> )	g-C <sub>3</sub> N <sub>4</sub> prepared from melamine powder, Cdots/3DGH prepared by dissolving glucose 10 mL. GO oxide, followed by Autoclave	A p E.coli, Ratiometric Photoelectrochemi cal	2.9 -2.9 $\times10^6$ CFU mL <sup>-1</sup>	0.66 CFU mL <sup>-1</sup>	Stability, good reproducibility, & repeatability		Milk		[119]	
CdSe quantum dots (QDs) sensitized $\text{Ho}^{3+}/\text{Yb}^{3+}\text{-TiO}_2$	CdSe QDs size 5nm, CdSe QDs increases the photocurrent response of TiO <sub>2</sub> & enabled it to be excited under the visible light	A b V.parahaemolyticus Photoelectrochemi cal	$10^1\text{-}10^9$ CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup>	Excellent selectivity, High sensitivity, RSD 5 electrode 1.2%		Shellfish products		[107]	
AuNPs integrated with ZnAgInS QDs	AuNPs <sub>C</sub> ZnAgInS QDs 10 nm, AuNPs integrated with ZnAgInS QDs with EDC/NHS with the help of mercaptoacetic acid, AuNPs <sub>C</sub> & ZnAgInS QDs ratio 1:100 is the best ratio	A b Hepatitis B virus surface antigen Photoelectrochemi cal	$5\times10^{-9}\text{-}3\times10^{-5}$ g L <sup>-1</sup>	$5\times10^{10}$ g L <sup>-1</sup>	Selective, GNP <sub>s</sub> /ZnAgInS/GCE showed 33 times higher photo current in comparison ZnAgInS/GCE, 94.5% of initial signal after 2 week		Human serum		[108]	
AuAg Nanoshells	AuAg NSs with increased resistance to chemical oxidation generates a strong electrochemical signal.	Escherichia coli & Salmonella	DP V	$10^2$ CFU mL <sup>-1</sup>	Low cost, Fast (10min), Easy to operation, need any additional reagent, substrate, AuAg NSs act as a natural redox enzymes		water		[120]	
	Monodisperse hollow AuAg NSs 60 nm with a thin outer shell of 10 nm,	Typhimurium			electrochemical signal					

photoelectrochemical aptasensor based on three-dimensional graphene hydrogel-loaded with carbon quantum dot (C-dots/3DGH) and graphene-like carbon nitride sheet (g-C<sub>3</sub>N<sub>4</sub>); the proposed sensor presented superior photoelectrochemical ability. Applying different bias voltage, the cathodic and anodic current produced, respectively, C-dots/3DGH and g-C<sub>3</sub>N<sub>4</sub> could be detected and differentiated; the

proposed dual signal system allowed not only to detect selectively an analyte but also to take into account possible influence of the environment on the sensor response. To demonstrate this, the authors reported on an *E.coli* aptasensor in which the aptamer was immobilized on the C-dots/3DGH surface. In the presence of *E.coli* if on one end the cathodic current, associated with the C-dots/3DGH, was reduced due to the



**Fig. 3.** A) Schematic of pyrolytic biosensor schema with permission ref [125]. B) Liposome enhanced plasmonic immunosensor with permission ref [126], C) Quantitative Immunoassay Based on SiO<sub>2</sub>@PAA@CAT-Catalyzed growth of AuNPs with permission ref [127].

blocking effect of aptamer and target on the other end no effect on the anodic current was recorded; subsequently variation in the anodic current was used to estimate environmental change. The quantitative measurement of the target pathogen was carried out by calculating the ratio of cathodic current to anodic current. The aptasensor showed a linear range of (2.9 CFU mL<sup>-1</sup> to  $2.9 \times 10^6$  CFU mL<sup>-1</sup>) with an excellent LOD of 0.66 CFU mL<sup>-1</sup> [119]. The schematic of this procedure is shown

in Fig. 2D

Simple and facile detection of *S.Typhimurium* and *E.coli* was demonstrated by the use of hollow electrocatalyst Au-Ag nanoshells that enabled the in situ generations of the readable electrochemical signal; *S.Typhimurium* and *E.coli* were mixed with PVP-coated Au-Ag nanoshells and incubated for short time. To generate an electrochemical signal the mixture was dropped to SPCE. Au-Ag nanoshells were deposited, by

**Table 2**

Colorimetric biosensors, List of Abbreviation Colorimetry (CO) [248–276].

Nanomaterial	Nanomaterial Remark	Bio	Pathogen	Metho d	Liner range	LOD	Remark	Real sample	Ref
(Magnetic nanobeads) MNP-COOH	<i>S. aureus</i> proteases peptide		<i>Methicillin-Resistant S.aureus</i>	Visual	7.5-7.5×10 <sup>6</sup> CFU mL <sup>-1</sup>	7CFU mL <sup>-1</sup>	40 and 100 CFU mL <sup>-1</sup> in real sample, Selective, 6-month stability	Ground beef, turkey sausage, Lettuce, milk	[122]
MNP-COOH	<i>E. coli</i> O157: H7 proteases peptide		<i>E.coli O157: H7</i>	Visual	121-1.21×10 <sup>6</sup> CFU mL <sup>-1</sup>	12 CFU mL <sup>-1</sup>	30-300 CFU mL <sup>-1</sup> in complex food matrices, rapid response time, 6 months, stability. Selective, Low cost and, Ease of use	Ground beef, turkey sausage, Lettuce, milk	[123]
MNP-COOH	<i>P. gingivalis</i> proteases peptide		<i>Porphyromonas gingivalis</i>	Visual	10 <sup>1</sup> -×10 <sup>9</sup> CFU mL <sup>-1</sup>	49CFU mL <sup>-1</sup>	Easy to perform, Fast (30 s response time)	Saliva samples	[124]
MNP-COOH	<i>Listeria</i> protease peptide		<i>Listeria monocytogenes</i>	Visual	2.17 x10 <sup>2</sup> -2.17 x10 <sup>8</sup> CFU mL <sup>-1</sup>	2.17 x10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, Easy to perform, Fast (30 s response time)	Whole milk Ready to eat meat	[125]
Blue, orange, and green dye coated polymer nano-beads	Ab Lactoferrin		<i>S. Typhimurium</i> , <i>S.enteritidis</i> , <i>S. aureus</i> and <i>C. jejuni</i> .	Visual	10-10 <sup>8</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective Portable, User friendly 10 CFU mL <sup>-1</sup> <i>S.Typhimurium</i> and <i>C.rjekuni</i> , 100 for <i>S. enteritidis</i> <i>S. aureus</i> CFU mL <sup>-1</sup> meat surface	Chicken meat	[248]
AuNPs <sub>c</sub> , Nanoliposome	Cysteine-loaded liposomes pH 7.25, AuNPs shows a very high stability	A b	<i>E. coli</i> O157: H7, <i>S.Typhimurium</i> <i>L.monocytogenes</i> .	Visual	10 <sup>1</sup> -10 <sup>3</sup> CFU mL <sup>-1</sup>	Single-digit pathogen	Enzyme free, <i>E.coli O157: H7</i> Milk, <i>S.Typhimurium</i> Apple juice, <i>L.monocytogenes</i> . Ground beef	[126]	
Blue Silica Np, Blue Silica Np-NH <sub>2</sub> , Blue Silica Np-COOH, MNPs- COOH	Blue Silica Np spherical shape 45nm, Reverse microemulsion method, tiny color change in basic solution& faintly lighter in acidic solution	A b	<i>S. pullorum</i> & <i>S. gallinarum</i>	Visual	8.8×10 <sup>1</sup> -8.8×10 <sup>9</sup> CFU mL <sup>-1</sup>	8.8×10 <sup>1</sup> CFU mL <sup>-1</sup>	Milk powder Inexpensive, Specific, Immune Np stable for 90 days	Milk powder [70]	
Blue SilicaNp, Blue Silica Np-NH, Magnetic bead	Blue Silica Np spherical shape 50nm, Reverse microemulsion method, color Stable in acidic &basic solution	A b	<i>S. pullorum</i> & <i>S. gallinarum</i>	Visual	10 <sup>1</sup> -10 <sup>5</sup> CFU mL <sup>-1</sup>	10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Non- enzymatic, Immune Np stable for 90 days	Chicken serum [71]	
AuNPs	AuNPs coated with streptavidin larger than 13 nm	A b	<i>E.coli</i> & <i>Salmonella</i>	Visual	10-10 <sup>3</sup> CFU mL <sup>-1</sup>	10CFU mL <sup>-1</sup>	Selective, Detect only live cell, total assay time is about 30 min when the sample is gently agitated	Lake water, Whole milk [249]	
AuNPs	AuNPs coated with streptavidin larger than 13 nm	A b	<i>Salmonella</i>	Visual	10-10 <sup>3</sup> CFU mL <sup>-1</sup>	10CFU mL <sup>-1</sup>	Selective	Tomato [250]	
SilicaNp MNP with SiO <sub>2</sub>	Silica Np 50 nm reverse emulsion, MNP Co-participation	A b	<i>S.pullorum</i> and <i>S. Gallinarum</i> .	Visual/CO	8.4 × 10 <sup>3</sup> 8.4 × 10 <sup>7</sup> CFU mL <sup>-1</sup>	1.7 × 10 <sup>3</sup> CFU mL <sup>-1</sup>	Specific, 96- 55% of the initial signal were preserved after storage of HRP-IgG-SiNps &IgG -MN P for 30- 120 days	Chicken liver [72]	
SiO <sub>2</sub> @PAA@C AT, MNP-COOH	SiO <sub>2</sub> +Polyacrylic acid size 80 nm Stöber method, Magnetic particle 180nm, High loading of catalase enzyme By SiO <sub>2</sub> nanoparticles modified with PAA brushes	A b	<i>Listeria monocytogenes</i>	Visual /pELISA	8 × 10 <sup>1</sup> -8 × 10 <sup>8</sup> CFU mL <sup>-1</sup>	8CFU mL <sup>-1</sup>	5 order of magnitude more sensitive than	Lettuce [127]	
AuNPs		A b	<i>Salmonella enterica</i>	Visual /CO	10 <sup>3</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	500 CFU mL <sup>-1</sup>	Selective,	Milk [251]	
MNP, AuNPs	Fe <sub>3</sub> O <sub>4</sub> - COOH 180 nm Dextran-capped AuNPs mono-dispersed particles with 15 nm	A b	<i>E. Coli</i> O157: H7	Visual CO	10 <sup>2</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	41 CFU mL <sup>-1</sup>	Selective	Milk [252]	
MNC/Pt	Fe <sub>3</sub> O <sub>4</sub> Hydrothermal method MNC/ Pt100nm,PtNps 16nm	A b	<i>E.coli O157: H7</i>	Visual/CO	10 <sup>1</sup> -10 <sup>5</sup> CFU mL <sup>-1</sup>	10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, MNcs/ <i>E. Coli</i> were separated from free MNcs by magnetophoretic Chromatography, Easy to perform, fast, Strong catalytic activity of Pt particle cause low LOD	Milk [136]	
AuNPs <sub>c</sub>	AuNPs probes labeled with HRP as probes to produce signal created by catalyzed TMB reaction	A b	<i>E.coli O157: H7</i>	CO	10 <sup>2</sup> -10 <sup>5</sup> CFU mL <sup>-1</sup>	100 CFU mL <sup>-1</sup>	Net fishing enrichment method, 200 CFU mL <sup>-1</sup> Pineapple juice	Pineapple juice [253]	
MNP-COOH	MNP-COOH 100nm coated with vancomycin	A L P- Ig G	<i>S. aureus</i>	CO	12-1.2×10 <sup>6</sup> CFU mL <sup>-1</sup>	3.3 CFU mL <sup>-1</sup>	Specific	Lake water, Milk, Urine Saliva [254]	
MNPs,AuNPs, Ag nanoclusters	MNPs Hydrothermal method AuNPs <sub>c</sub> spherical 18.1nm	A b/ A p	<i>Listeria monocytogenes</i>	Visual/ CO	10-10 <sup>6</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Specific	Pork [255]	
Au-CNT nanohybrids	Au CNT nanohybrids in situ synthesis with HCOONa The optimum 10 nm	A b	<i>Influenza virus A (H3N2)</i>	Visual /CO	10-50000 PFU mL <sup>-1</sup>	3.4 PFU mL <sup>-1</sup>	100 times and 500 times more sensitive than ELISA& commercial kit	[135]	
AuNPs	AuNPs 13nm	A b	<i>Influenza A virus H3N2</i>	Visual/CO	0-80 HAU	7.8 HAU	Specific, Easy to operate	[256]	
Magnetic Nano beads, AuNPs	Superparamagnetic Nano beads 200nm Electrodeposited AuNPs	A b	<i>Human enterovirus 71 (EV 71)</i>	Visual	1× 10 <sup>11</sup> -6×10 <sup>7</sup> g mL <sup>-1</sup>	1× 10 <sup>9</sup> Visual g mL <sup>-1</sup>	Selective, Investigate two different method, No pre-treatment or solution transfer,	Clinical serum sample [257]	

AuNPs	AuNPsc 20 nm for a capture probe, cationic cystamine 40nm, CTAB capped AuNPs 30 nm core/shell-like structure, pr cystamine-AuNPs capable of ob acting as hydrogen bond and enhance color detection	R N	<i>Hepatitis C virus RNA</i>	Visual/CO	1-3 IU $\mu\text{l}^{-1}$	4.57 IU $\mu\text{l}^{-1}$	Simple, Rapid, RNA folding impact on the aggregation behavior	Clinical serum sample	[128]
AuNPs film AuNPs + CdTe QDs	Coating AuNPs on glass, 96-well polystyrene plates, Au NPs more densely deposited onto the hydrophobic surface than onto the hydrophilic surface, CdTe QDs 2.85 nm, +AuNPs 40 nm prepared with cystamine solution	A b &	<i>Influenza virus A H1N1 H3N2</i>	Visual/CO	$10^{-8}\text{ g L}^{-1}$ - $10^{-2}\text{ g L}^{-1}$ $10^{-50,000}$ PFUA $\text{mL}^{-1}$	$5.05 \times 10^{11}\text{ g mL}^{-1}$ 24.3 PFUA $\text{mL}^{-1}$ PFUA $\text{mL}^{-1}$	The peroxidase-like catalytic activity of Au NP films and colloidal AuNPs, 116 times better than ELISA		[130]
AuNPs <sub>c</sub> , Magnetic nanobeads QD	Superparamagnetic 500nm, In situ, synthesized QDs with S.aureus,	A b	<i>Enterovirus 71</i>	Visual/CO	$5 \times 10^{-7}$ - $3 \times 10^{-6}$ $\text{g L}^{-1}$		Selective, nanoparticles,	Self-synthesized	Human throat swab [258]
AuNPs	AuNPsc p	A p	<i>C. sakazakii</i>	Visual	$7.1 \times 10^3$ $7.1 \times 10^7$ CFU $\text{mL}^{-1}$	$7.1 \times 10^3$ CFU $\text{mL}^{-1}$	Selective, Easy to operate	Powdered infant formula	[259]
AuNPs	AuNPsc p	A p	<i>Campylobacter coli</i> & <i>Campylobacter jejuni</i>	Visual	$10^7$ - $10^8$ CFU $\text{mL}^{-1}$	$7.2 \times 10^3$ CFU $\text{mL}^{-1}$	Can distinguish live and heat-killed bacteria,	Chicken carcass	[260]
Au@Pd nanoparticles	The thickness of palladium layer 1-2 nm cause a redshift for Au@PdNps	A p	<i>Campylobacter jejuni</i>	Visual/ CO	$10^{-10}$ CFU $\text{mL}^{-1}$	$100$ CFU $\cdot \text{mL}^{-1}$	Selective, After 6 months RSD peroxidase-like activity change 4.2%,	Milk	[40]
AuNPs	AuNPsc peroxidase-like activity of AuNP17 nm	A p	<i>Pseudomonas aeruginosa</i>	CO/electrochemical	$60.0$ to $6.0$ $\times 10^7$ CFU $\text{mL}^{-1}$	$60.0$ CFU $\text{mL}^{-1}$	Selective, Applying two different method	-	[261]
Au Nanorods, Magnetic nanoparticle, MnO <sub>2</sub> Nanoparticle	Au Nanorods silver ion-assisted seed-mediated growth a monodisperse and uniform rod.Magnetic nanoparticle 242.7 nm	A p/ A b	<i>Listeria monocytogenes</i>	Visual/CO	$10^{-5}$ CFU $\text{mL}^{-1}$	$10$ CFU $\text{mL}^{-1}$	Selective, Detection in pork sample Without pre-enrichment, MnO <sub>2</sub> peroxidases like nano artificial enzyme	Minced pork	[133]
ZnFe <sub>2</sub> O <sub>4</sub> /rGO	ZnFe <sub>2</sub> O <sub>4</sub> /rGO Good peroxidase-activity, Stable than HRP	A p	<i>S.Typhimurium</i>	Visual/CO	$11$ - $1.10 \times 10^5$ CFU $\text{mL}^{-1}$	$11$ CFU $\text{mL}^{-1}$	Selective	Milk	[134]
Fe <sub>3</sub> O <sub>4</sub> MNP	MNP 20nm co-participation	A p	<i>S. Typhimurium</i>	Visual/CO	-	$7.5 \times 10^5$ CFU $\text{mL}^{-1}$	Peroxidase behavior Fe <sub>3</sub> O <sub>4</sub> MNP	-	[131]
Fe <sub>3</sub> O <sub>4</sub> Nano cluster	Fe <sub>3</sub> O <sub>4</sub> Solvothermal method Fe <sub>3</sub> O <sub>4</sub> Nano cluster Fe <sub>3</sub> O <sub>4</sub> + poly-L-lysine layer Show higher catalytic activity	A p	<i>Listeria monocytogenes</i>	Visual/CO	$5.4 \times 10^3$ - $\times 10^8$ CFU $\text{mL}^{-1}$	$5.4 \times 10^3$	Selective	Milk	[262]
AuNPs	AuNPsC 16 nm	A p	<i>S. Typhimurium</i>	Visual/CO	$10^7$ - $10^9$ CL $10$ - $10^9$ AM CFU $\text{mL}^{-1}$	$10$ CL, $100$ Am CFU $\text{mL}^{-1}$	Sensitive, Peroxidase behavior of AuNPs	Apple juice	[132]
AuNPs	AuNPsc p	A p	<i>Sh. flexneri</i>	Visual/CO	$10^2$ - $6$ CFU $\text{mL}^{-1}$	$80$ CFU $\cdot \text{mL}^{-1}$	Selective, Portable	Smoked salmon	[263]
AuNPs	AuNPsc p	A p	<i>E. coli O111:B4LPS</i>	Visual/CO	$2.5$ - $20 \times 10^3$ g $L^{-1}$	$1 \times 10^3$ g $L^{-1}$	Specific		[264]
Au@Pt NR	NRs aspect ratio 3.8, Pt nanodots with sizes of 3-4 nm	A b	<i>Measles IgM antibodies</i>	Visual / LAMP	$10^{-5}$ - $10^{-3}$ g $L^{-1}$	$10^{-3}$ g $L^{-1}$	1000 times more sensitive than ELISA, More stable in different temperatures and pH than HRP	-	[265]
AuNPs	AgNPsc 20nm		Phosphatidylcholine-phospholipase C gene (plcB) <i>Listeria monocytogenes</i>	Visual / LAMP	$2.82$ - $2.82 \times 10^8$ CFU $\text{mL}^{-1}$	$8 \times 10^{10}$ g and CFU $\text{mL}^{-1}$	Fast, Specific, 10 times more sensitive than the conventional PCR assay	Chicken Meat	[266]
AuNPs	AuNPsc 10 to 20 nm		DNA probe <i>Hepatitis C Virus (HCV) RNA</i>	Visual/CO	$3.70 \times 10^7$ - $3 \times 10^6$ mol $\text{L}^{-1}$	$5 \times 10^{-7}$ mol $\text{L}^{-1}$	Investigate free patchy end in differential adsorption of ssDNA and dsDNA over citrate-modified Au NPs and set up rules	-	[267]
AuNPs	AuNPsc 13nm		Genomic DNA from bacterial cultures <i>Emetic Bacillus cereus</i>	CO/asym metric PCR	$9.2 \times 10^1$ $9.2 \times 10^6$ CFU $\text{mL}^{-1}$	$9.2 \times 10^1$ Milk $9.2 \times 10^2$ CFU $\text{mL}^{-1}$	Selective, Viable detection of the pathogen	Milk	[268]
AuNPs	AuNPsc 12.867 ± 2.399 nm and 13.705 ± 2.557 nm		DNA probe <i>Mycobacterium avium DNA</i>	Visual		$1.03 \times 10^{-2}$ g $\text{L}^{-1}$	Specific	-	[269]
Flower-shaped AuNPs	Flower-shaped AuNPs 68.7 ± 2.8 nm, AuNPs + planet extract		DNA probe <i>Amplicons of hlyA gene &amp; genomic DNA of Listeria monocytogenes</i>	Visual	$4.84 \times 10^{-8}$ $1.004 \times 10^{-7}$ g	$4.84 \times 10^{-8}$ g	Green synthesis method, Fast, Simple, Selective	-	[270]
AuNPs	AuNPs 6-10 nm		DNA probe <i>DNA S. aureus</i> ,	Visual/CO	$5$ - $40 \times 10^3$ g $\text{L}^{-1}$	$8.37 \times 10^3$ g $\text{L}^{-1}$	Selective	-	[271]
CTAB capped gold nanoshells	CTAB capped gold nanoshells 23±0.58	-	<i>S.aureus, P.aeruginosa, S. enterica, E. coli</i>	Visual/CO	$10^1$ - $10^7$ cells $\text{mL}^{-1}$	$10$ CFU $\text{mL}^{-1}$	Rapid, Simple, Without biomolecule, Answer to a broad range of pathogen	-	[272]

AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 18±3 nm	-	<i>E.coli</i>	Visual	$10^4\text{--}10^7$ CFU mL <sup>-1</sup>	$1.02 \times 10^3$ CFU mL <sup>-1</sup>	Rapid, reliable, and cost-effective, Without biomolecule	-	[273]
Au nanorods	Au nanorode seed-mediated growth method. 59 ± 8 nm and a width of 11 ± 1 nm	-	<i>E.coli</i>	Visual/ CO	-	$1 \times 10^5$ CFU mL <sup>-1</sup>	Simple, Rapid inexpensive, portable, disposable devices, User friendly, without biomolecule	-	[274]
AuNPs	AuNPs <sub>C</sub> 20 nm	-	<i>E. coli, S.Typhimurium, Klebsiella pneumonia</i>	Visual, FRET	$10\text{--}10^7$ cells mL <sup>-1</sup>	$10$ cells mL <sup>-1</sup>	Rapid, Simple, Without biomolecule, Answer to a broad range of pathogen	Tap water and lake water	[129]
Van-coated Au NPs	9.5 nm, Molar ration 1:1 HAuCl <sub>4</sub> to Van the most stable	-	Gram-positive bacteria	Visual	$0\text{--}8 \times 10^8$ CFU mL <sup>-1</sup>	$1 \times 10^9$ CFU mL <sup>-1</sup>	Ag enhancement, Without biomolecule	Orange juice Tap water	[275]
MPBA-AgNPs	AuNPs citrate & NaBH <sub>4</sub>	-	Gram-negative bacteria	Visual/U V/Smart phone	$5 \times 10^4\text{--}1 \times 10^7$ CFU m L <sup>-1</sup>	$9 \times 10^3$ CFU m L <sup>-1</sup>	Rapid, Low cost, Selective Without biomolecule		[276]

applying a constant potential followed by a DPV scan to oxide the residual Ag atoms to Ag<sup>+</sup> in PBS solution. The proposed electrode was demonstrated to be able to measure  $10^2$  CFU mL<sup>-1</sup> of bacteria without using specific biomolecule [120]. More examples of the electrochemical bio/sensors are presented in Table 1.

### 3.2. Colorimetric biosensors

Colorimetry is a well-established method with widespread use in sensor development; this transduction method presents few advantages as low cost, simple/no instrumentation, and ease of fabrication. Naked-eye detection is a fascinating branch of colorimetric methods that enables qualitative detection of analytes; on the other end, the use of spectrophotometric reading allows quantitative measurements [121].

Enzyme induced disaggregation of black magnetic nanobeads (MB) network, based on peptidic crosslinking, was explored by the Zourob team, in the development of optical sensing of several pathogens. MB networks were placed on gold substrates, located on a paper surface, and kept in position with a magnet. Upon addition of bacterial extract, that contains proteases to the sensing surface, disaggregation of the MB network, due to the peptide digestion, take place exposing in this way the underneath Au. The percentage of the observable area was shown to be proportional to the bacteria protease concentration. The proposed sensors could achieve LOD of 7, 12, 49,  $2.17 \times 10^2$  CFU mL<sup>-1</sup> for *Methicillin-Resistant S.aureus*, *E.coli O157: H7*, *Porphyromonas gingivalis*, *Listeria monocytogenes* respectively [122–125]. The schematic of this procedure is presented in Fig. 3A. Nanoliposomes contain cysteine were demonstrated in the development of optical ELISA tests for the detection of *Salmonella*, *Listeria*, *E. coli O157*, and rabbit IgG antibody. In the proposed strategies the nanoliposomes were used as a label in the sandwich assay. Optical detection was made possible by the ability of the cysteine, released following rupture of the nanoliposome, to aggregate AuNPs with a subsequent blue shift of the AuNPs plasmonic. The purposed sensor showed 6 orders of magnitude enhancements for IgG detection, down to aM levels, in comparison to the conventional ELISA method which used HRP as a biocatalyst for TMB reduction [126]. The schematic of this assay was illustrated in Fig. 3B.

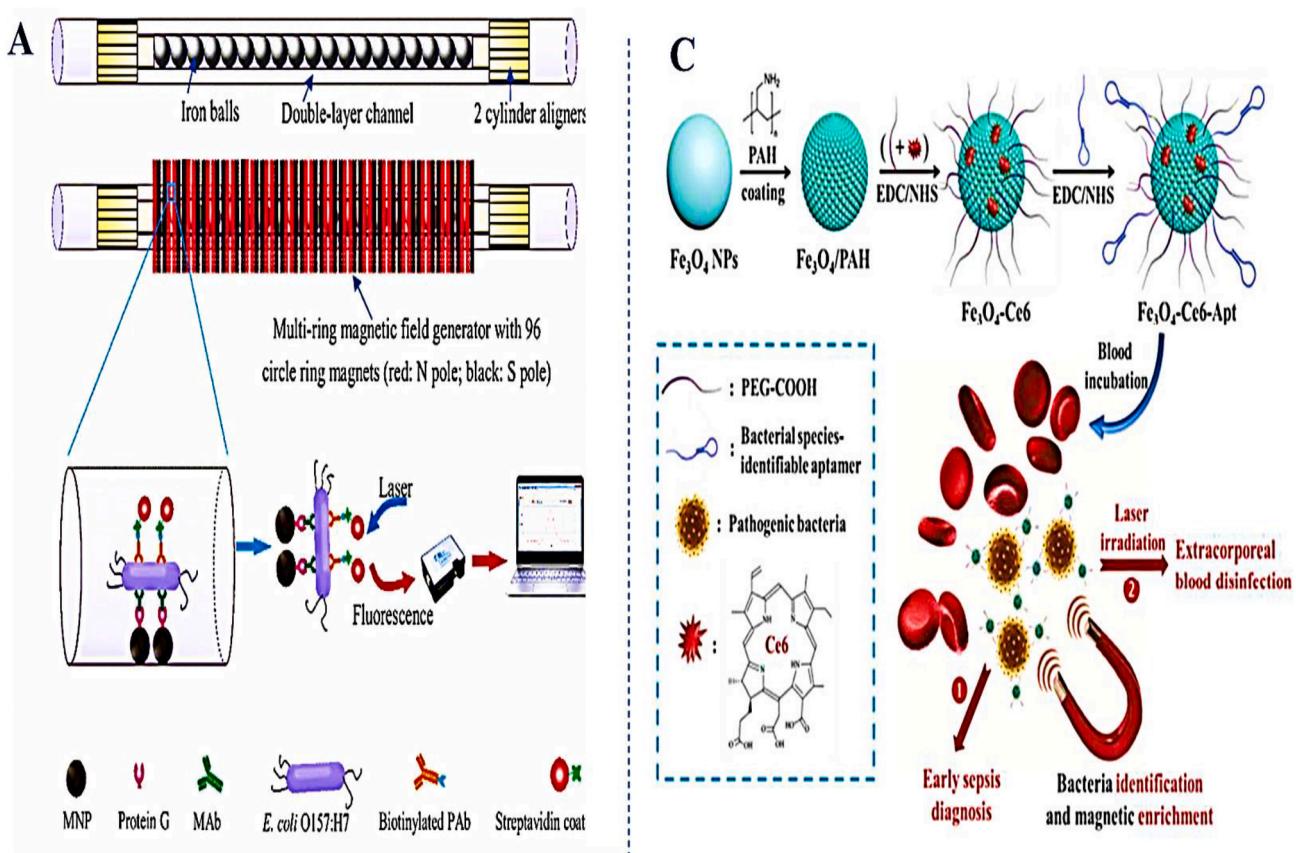
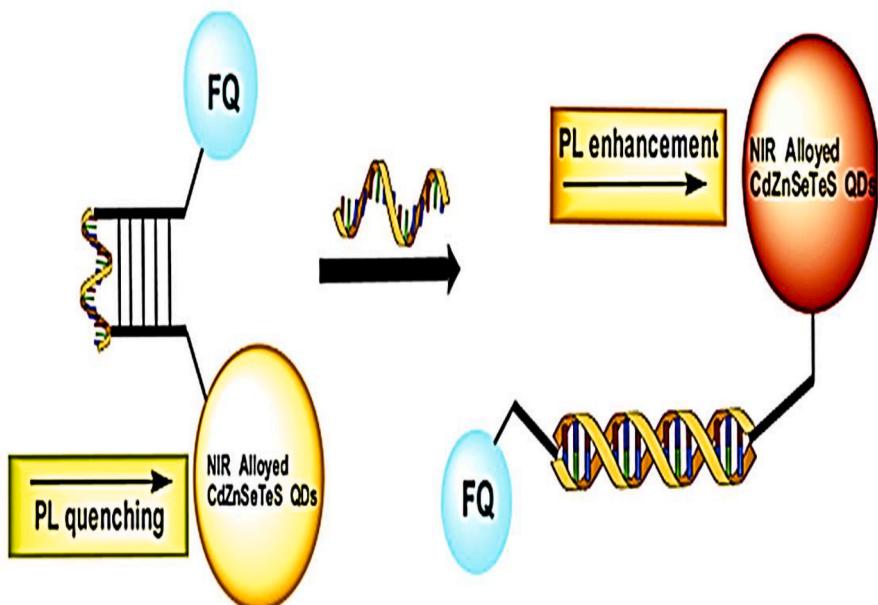
Silica nanoparticle with poly(acrylic acid) brushes worked as a carrier for high loading of catalase enzyme; the enzyme loaded particles were used as a label in ELISA for sensitive visual detection of *L.monocytogenes* ( $10$  CFU mL<sup>-1</sup>). The proposed assay was based on the differences in AuNPs formation when different concentrations of the reducing agent (hydrogen peroxide) were present in the reaction chamber. In the absence of target analyte the fast reduction of gold ions by hydrogen peroxide, both added in the ELISA well, quasi-spherical non-aggregated gold nanoparticles with red color were formed. By contrast in the presence of analyte the catalase enzymes, loaded on the silica labels, were then able to degrade the hydrogen peroxide giving rise to an aggregated nanoparticle with characteristic blue color. High loading of the enzyme increased LOD 2 and 5 orders of magnitude in comparison with catalase plasmonic ELISA (without silica nanoparticle) and HRP

based ELISA methods [127]. The schematic of this assay is depicted in Fig. 3C.

Gold nanomaterials are among the most used material in colorimetric detection. The different detection approached based on Au nanomaterials can be classified into three “groups”: (i) approaches based on aggregation/disaggregation, (ii) approaches based on catalytic properties of the Au nanomaterials, and (iii) growth of hybrid nanomaterials [9]. AuNPs modified with a thiolated Hepatitis C Virus (HCV) specific probe was used to detect unamplified HCV RNA. In the proposed assay probe modified AuNPs were incubated with unamplified HCV RNA at 95 °C for a fixed time; two positively charged gold nanoparticles (cysteamine and CTAB) capped used to promote aggregation of the above-mentioned AuNPs. When no complementary target was present in the solution aggregation of the AuNPs, due to electrostatic interaction was recorded (red to blue color change); on the other end in the presence of HCV RNA the solution remained red. Cysteamine capped AuNPs provided sharper color. The biosensor showed a LOD of  $4.57 \text{ IU} \mu\text{L}^{-1}$  [128]. AuNPs were also used to transduce the interaction of antibiotics with the bacterial outer membrane; this was shown to enable the sensitive detection of bacteria down to  $10\text{CFU mL}^{-1}$ . More specifically in their proof of concept work, Singh et al. used Colistin, a cationic antibiotic, as a bacteria recognition element and as an AuNPs aggregation inducing element [129].

Visual detection can be enhanced by the catalytic activity of nanomaterials to a chromogenic substrate like 3,3',5,5'-tetramethylbenzidine (TMB) [130–132]. Peroxidase activity of nanomaterial has been also exploited in colorimetric sensing. For example, a sandwich-type biosensor based on apt-MNP and IgY- BSA-MnO<sub>2</sub> nanoparticles was reported for the detection of *L. monocytogenes*. The MnO<sub>2</sub> in the label was then used to oxidize TMB to TMB<sup>2+</sup>; oxidizedTMB<sup>2+</sup> reacted with AuNRs resulting in a gradual degradation of the AuNRs with associated variation in their aspect ratio. The shift of longitudinal localized surface plasmon was shown to correlate linearly with the log concentration of *L. monocytogenes* in the  $10\text{--}10^6$  CFU mL<sup>-1</sup> concentration range [133]. Wu et al. developed a colorimetric sandwich-type aptasensor in which ZnFe<sub>2</sub>O<sub>4</sub>/reduced GO nanocomposite was used to catalyze the oxidation of the TMB by consuming H<sub>2</sub>O<sub>2</sub> that resulted in the formation of a blue color product which is then measured by micro-reader; detection of *Salmonella enterica serovar* down to  $11$  CFU mL<sup>-1</sup> was demonstrated [134]. In another study, the nanocomposite of AuNPs and CNT was shown to have higher catalytic activity in comparison to AuNPs and CNT alone toward the oxidation of TMB by H<sub>2</sub>O<sub>2</sub>. The proposed nanocomposites were utilized in an immunosensor for the sensitive detection of influenza A H<sub>3</sub>N<sub>2</sub> virus down to  $10$  PFU mL<sup>-1</sup> [135].

Magnetic nanoparticles covered with platinum (Pt/MNC) were exploited as a dual functional element in an *E.coli O157: H7* immunoassay; these were used to (i) carry the biorecognition element, half-fragment of monoclonal *E.coli O157: H7* antibody, and as (ii) catalyst for the oxidation of TMB. Nanoparticles were added to milk solution containing *E.coli O157: H7* followed by incubation for short time and separated with a permanent magnet. A precision pipette was used to

**B**

**Fig. 4.** A)The illustration of double channel coated by immunomagnetic bead and QDs with permission ref [139], B) The Schematic of NIR alloyed QD-MB designed biosensor with the permission of ref [144] C) The pattern of the  $\text{Fe}_3\text{O}_4\text{-Ce6-Apt}$  system for early sepsis diagnosis [148].

**Table 3**

Fluorometric biosensors Abbreviation Fluorescence (FL), Fluorescence energy transfer resonance (FRET) Emission (em), Reverse assay strategy (RAS) [277-312].

Table 3

Nanomaterial	Nanomaterial remark	Bi o	Pathogen	Method	Linear range	LOD	Remark	Real Samp le	Ref
Magnetic nanobeads-COOH CdTe QD MPA	Aqueous synthesis of QD, Fluorescence of mercaptopropionic acid-modified CdTe QDs is highly sensitive to H <sub>2</sub> O <sub>2</sub> , λ <sub>abs</sub> =566 nm, λ <sub>em</sub> =590 nm, Carboxylated magnetic nanoparticles 180 nm	A b	<i>E. coli</i> O157: H7	FL/Quenching	1.18×10 <sup>3</sup> 1.18×10 <sup>6</sup> CFU mL <sup>-1</sup>	5×10 <sup>2</sup> CFU mL <sup>-1</sup>	Specific, Novel method based on extreme bioactivity of catalase to H <sub>2</sub> O <sub>2</sub> & sensitive H <sub>2</sub> O <sub>2</sub> QD	Milk	[277]
MNP QD	Carboxylated magnetic nanoparticles 180 nm CdSe/ZnS core shell, λ <sub>em</sub> =614 nm	A b	<i>E. coli</i> O157: H7	FL	8.9×10 <sup>0</sup> 8.9×10 <sup>5</sup> CFU mL <sup>-1</sup>	1.4×10 <sup>1</sup> CFU mL <sup>-1</sup>	Specific	Milk	[139]
Fe <sub>3</sub> O <sub>4</sub> @Au CdTe QD	CdTe QD 3nm, Chitosan Coated CdTe 40 nm Fe <sub>3</sub> O <sub>4</sub> @Au, Fe <sub>3</sub> O <sub>4</sub> Co-precipitation Au Shell CTAB	A b	<i>E. coli</i>	FL	10 <sup>2</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	30 CFU mL <sup>-1</sup>	Selective, Fast, Sensitive	Urine	[278]
QD, Magnetic beads	Carboxyl QDs 2–10 nm in size Magnetic beads (1150 nm)	A b	<i>E. coli</i> O157: H7 & <i>Enterica</i>	FL/RAS	S. enterica 10 <sup>2</sup> -10 <sup>7</sup> <i>E. coli</i> 50-10 <sup>8</sup> CFU mL <sup>-1</sup>	E.coli 30 CFU mL <sup>-1</sup> S. enterica 60 CFU mL <sup>-1</sup>	Multiplex, 50 time sensitivity enhancement than ELISA	Milk	[279]
CdSe /ZnS AuNPs	core-shell CdSe/ZnS QDs AuNPs citrate 20nm	A b	<i>Vibrio parahaemolyticus</i>	FL Turn on	10-10 <sup>4</sup> CFU mL <sup>-1</sup>	1050CFU mL <sup>-1</sup> in real sample 50CFU mL <sup>-1</sup>	Selective, core-shell CdSe/ZnS QDs stable for 2 month, Shrimp	[142]	
CdSe/ZnS QD	CdSe/ZnS QD 5-8 nm	A b	<i>Salmonella</i>	Visual/F L	-	10cell mL <sup>-1</sup>	Selective, Simple, POC analysis.	Stool	[280]
MNP-COOH liposomes	MNP-COOH 30 nm Liposomes 244.3nm	A b	<i>C. sakazakii</i>	FL	10 <sup>2</sup> -10 <sup>7</sup> CFU ml <sup>-1</sup>	3.3×10 <sup>3</sup> CFU ml <sup>-1</sup>	2 cells/10g After 6 h enrichment 10 <sup>4</sup> CFU mL <sup>-1</sup> without pre-incubation	Powdered infant milk	[281]
CdTe-COOH AuNPs 14.5	two particles become less than 10 nm and AuNPs citrate 14.5 nm+ 11-MUA	A b	<i>Vibrio choleraeouter membrane protein W</i>	FL/FRE T	2 –10 × 10 <sup>-9</sup> mol L <sup>-1</sup>	2 × 10 <sup>-9</sup> mol L <sup>-1</sup>	Simple, Fast	-	[282]
AuMNPs CdSelNs QD	AuMNPs decorated with GO MNP Co-precipitation method	A b	Influenza virus H <sub>1</sub> N <sub>1</sub>	FL	1×10 <sup>12</sup> g L <sup>-1</sup> 1×10 <sup>9</sup> g L <sup>-1</sup>	7.27×10 <sup>12</sup> g L <sup>-1</sup> Human serum 6.07×10 <sup>9</sup> g L <sup>-1</sup>	Specific	Human serum	[143]
Alloyed quaternary CdSeTeS QDs AuNPs	Alloyed quaternary 3.8 nm CdSeTeS hot-injection organometallic route & capped with L-cysteine AuNPs 40 nm synthesized from HAuCl <sub>4</sub> in HEPES buffer with cysteine capping	A b	Influenza virus H <sub>1</sub> N <sub>1</sub>	FL	10 <sup>8</sup> - 10 <sup>7</sup> g L <sup>-1</sup>	3×10 <sup>11</sup> g L <sup>-1</sup> Human Serum 4×10 <sup>10</sup> g L <sup>-1</sup> 10 PFU mL <sup>-1</sup>	Selective	Human Serum	[283]
SiQDs AuNRs	Silica-coated quantum dots (SiQDs) QD 3.5 nm average size, Silica shell of 15 nm, the Aspects ratio of the AuNRs rise 3 times by increasing concentration of AgNO <sub>3</sub> , CTAB-coating over AuNRs worked as a bilayer, preventing aggregation and allowing its growth by face-sensitive surface adsorption	A b	Mycobacterium tuberculosis Ag85B antigen	FL	1×10 <sup>6</sup> -1×10 <sup>7</sup> g L <sup>-1</sup>	1.3×10 <sup>10</sup> gL <sup>-1</sup>	Selective	Urine	[284]
CdZnSeTeS QDs	QDs 9nm highly, Homogenous and monodispersed PL enhancement was considerably higher for the NIR alloyed QD-MB than for the conventional CdSe/ZnS QD-MB probe		molecular beacon strains of influenza virus H1N1 RNA	FL	2 -10 copies/mL	Specific		Human serum	[144]
AuNPs	AuNPsC		Ab/DNA probe <i>E. coli</i> O157:H7, <i>S. choleraesuis</i> <i>L. monocytogenes</i>	FL	10 <sup>1</sup> -10 <sup>7</sup> CFU m L <sup>-1</sup>	<i>E. coli</i> O157: H7 3.4 × 10 <sup>1</sup> , <i>S. choleraesuis</i> 6.4 × 10 <sup>0</sup> , <i>L. monocytogene</i> 7.0 × 10 <sup>1</sup> CFU m L <sup>-1</sup>	Selective, Simultaneous	Milk	[285]
FITC-doped Silica NPs SilicaNPs-COOH	Silica reverse microemulsion method monodisperse particles 60 nm FITC-doped silica NPs increase photostability of dye-doped silica	A b	<i>E. coli</i> O157: H7	FL microscopy & flow cytometry	9×10 <sup>5</sup> CFU mL <sup>-1</sup>	FITC-doped silica NPs brighter and higher photostability	Ground Beef	[286]	
UCNPs	Hexagonal uniform UCNP 50 nm	A b	<i>E. coli</i>	FL	42 to 4.2 × 10 <sup>7</sup> CFU mL <sup>-1</sup>	42 CFU mL <sup>-1</sup>	Selective	Pork Milk	[287]
UCNPs MNP	UCNP hexagonal shape 105 nm MNP hydrothermal method 110nm	A p	<i>E. coli</i>	FL	58 to 5.8 × 10 <sup>6</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective	Pork sample	[150]
NaYF <sub>4</sub> :Yb/r AuNPs	AuNPs monodispersed 13 nm NaYF <sub>4</sub> :Yb/Er monodispersed with an average diameter of 35 nm	A p	<i>Escherichia coli</i> 8739	FL/ FRET	5-10 <sup>6</sup> CFU mL <sup>-1</sup>	3 CFU mL <sup>-1</sup>	Specific	Tap/Pond water Milk	[145]
CdTe QD Fe <sub>3</sub> O <sub>4</sub>	CdTe CD 2nm	A p	<i>S. Typhimurium</i>	FL	10 <sup>1</sup> -10 <sup>10</sup> CFU m L <sup>-1</sup>	1CFU m L <sup>-1</sup>	Specific,	Water Milk	[140]

QD, Thulium-doped UCNPs, Magnetic bead	QD1-2 nm size Down-conversion of QDs & Upconversion of UCNP afford a suitable detection platform without spectral overlap	A p	<i>S. aureus</i> , & <i>S.Typhimurium</i>	FL	50-10 <sup>6</sup> CFU mL <sup>-1</sup>	16 and 28 CFU mL <sup>-1</sup> for <i>S.aureus</i> , & <i>S. Typhimurium</i>	Selective, detection, of <i>S.aureus</i> , & <i>S. Typhimurium</i>	Simultaneous	[141]
GO		A p	<i>S. enteritidis</i>	FL	10 <sup>2</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup>	Selective, truncated aptamer lower dissociation constant and 2 times lower LOD than full sequence	Milk	[288]
CDs , GO	CD	A p	<i>P. aeruginosa</i>	FL/FRE T	10 <sup>1</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	9CFU mL <sup>-1</sup>	Selective, Easy to perform, High- throughput sequencing method to select aptamer	Mineral water	[149]
Fe <sub>3</sub> O <sub>4</sub>	Fe <sub>3</sub> O <sub>4</sub> Co-participation method coated with silica layer with NH <sub>2</sub> group 283 nm, MCM-41 type silica particles 187 nm wit	Ap/DNA probe <i>S. aureus</i>	FL	800-10 <sup>4</sup> CFU mL <sup>-1</sup>	682 CFU mL <sup>-1</sup>	Selective	Whole blood	[289]	
MBs-COOH Van-BSA-MBs	MBs-COOH 180nm	A b	<i>S. aureus</i>	Flow cytomet ry	3.3×10 <sup>1</sup> 3.3×10 <sup>3</sup> CFU mL <sup>-1</sup>	- 3.3×10 <sup>1</sup> CFU mL <sup>-1</sup>	Selective, Sensitives Water, Milk, Fruit juice		[290]
Hollow silica nanosphere, Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> @ PAA	Hollow silica nanospheres 350 nm prepared by ammonia-catalyzed hydrolysis and condensation of TEOS in an aqueous ethanol solution. Fe <sub>3</sub> O <sub>4</sub> angular and amorphous particles with sizes ranging from 7 to 20 nm.,Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> quasi- spherical 30 to 50 nm	A b	<i>E. coli O157: H7</i>	FL	4-4×10 <sup>8</sup> CFU mL <sup>-1</sup>	3 CFU mL <sup>-1</sup>	Specific, bio-conjugated FHSNs stable for 60 days, 96.7% of initial response remained after 60 days	Milk, Orange juice, River water,	[146]
CdTe QD	CdTe QD 4.5nm	A p	<i>E. coli O157:H7</i>	FL	10 <sup>2</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, Recoveries 76-91%.	Meat, Green sprouts, Chicken Milk	[3]
AuNCs@Van	Vancomycin as reducing & stabilizing agent AuNCs uniform spherical and monodisperse with an AuNCs@Van diameter of 2.0±0.6 nm. Fluorescence stable in different buffers and at varying pH, only falling off significantly at pH below 4. The fluorescence quantum yield 13%	A p	<i>S. aureus</i>	FL	32-10 <sup>8</sup> CFU mL <sup>-1</sup>	70 CFU mL <sup>-1</sup>	Specific, Good sensitivity, Excellent linear range Recoveries 96.94% - 101.24%	Milk, Human serum	[291]
AuNCs@Van AuNPs	AuNCs@Van , Vancomycin as reducing & stabilizing agent	A p	<i>S. aureus</i>	FL	20-10 <sup>8</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective, Facile	Milk, Orange juice, Serum	[147]

CdSe/ZnS	Water-soluble QD	A p	<i>E. coli O157: H7</i>	FL	10 <sup>7</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Specific, Sensitive	Tap water	[292]	
QD	Red, yellow, green QDs	A P	<i>E. coli O157: H7</i>	FL	10 <sup>2</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, Long term stability, POC detection, In situ synthesis of photoluminescence-quenching nano paper	Lake water	[293]	
Fe <sub>3</sub> O <sub>4</sub> NaYF <sub>4</sub> : Ce/Tb NaGdF <sub>4</sub> : Eu	One-step solvothermal method synthesis, monodisperse, uniform, roughly spherical with 30 nm.	A p	<i>S. Typhimurium</i> & <i>S. aureus</i>	FL	10 <sup>7</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	15, 20 CFU mL <sup>-1</sup>	<i>S. Typhimurium</i> and <i>S. aureus</i>	Milk	[294]	
NaYF <sub>4</sub> : Ce/Tb	NaYF <sub>4</sub> : Ce/Tb 30 nm Solvothermal method 2- aminoethyl dihydrogen phosphate as a surfactant and capping agent.	A p	<i>S.Typhimurium</i>	TR- FRET	10 <sup>2</sup> -10 <sup>3</sup> CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup>	Selective, Good sensitivity	Chicken , Egg	[295]	
AuNPs <sub>c</sub> GQD	AuNPs 15nm acceptor of FRET pair GQD 3nm donor of FRET pair dispersed fine in water solution because of the abundant oxygen-containing functional	A p	<i>MecA</i> sequence of <i>S. aureus</i>	FL/FRE T	1×10 <sup>9</sup> -1×10 <sup>7</sup> mol L <sup>-1</sup>	1×10 <sup>9</sup> mol L <sup>-1</sup>	Selective, Sensitive PCR amplified product of <i>S. aureus</i> <i>mecA</i> gene		[296]	
Ag@SiO <sub>2</sub>	AgNPs 50 nm with 30nm silica shell (optimum thickness)	A p	Recombinant hemagglutinin (rHA) protein H5N1 influenza virus	FL/MEF	2×10 <sup>6</sup> -1×10 <sup>4</sup> g L <sup>-1</sup>	2×10 <sup>6</sup> g Human serum3.5×10 <sup>6</sup>	L <sup>-1</sup>	Selective, Fast, Ag@ SiO <sub>2</sub> increase the sensitivity 50 times, Recovery 105%	Human serum	[297]
Fe <sub>3</sub> O <sub>4</sub> -Ce6-Apt	Magnetic Fe <sub>3</sub> O <sub>4</sub> nanoparticles were synthesized by thermal decomposition of the iron-oleate complex. Compared to micro-sized magnetic particles the nano-sized Fe <sub>3</sub> O <sub>4</sub> -Ce6-Apt has a higher surface to volume ratio exhibits a higher binding capacity to targets.Fe <sub>3</sub> O <sub>4</sub> -Ce6-Apt nanosystem	A p	<i>S. aureus</i>	fluoresc ence microsc opic	-	10 CFU mL <sup>-1</sup>	Early sepsis diagnosis in mice (1.5h), Selective, Very good sensitivity, Fe <sub>3</sub> O <sub>4</sub> -Ce6-Apt 6 days of storage in blood, .No clear Potential toxicity of Fe <sub>3</sub> O <sub>4</sub> - Ce6-Apt toward red blood cells	Blood mice model	[148]	
MN <sup>2+</sup> NaYF <sub>4</sub> :Y b, Tm UCNP Au Nanorode	NaYF <sub>4</sub> : Yb, Tm UCNP spherical shape donor Au Nanorods Seed growth method acceptor	A p	<i>S.Typhimurium</i>	LET	12-5×10 <sup>5</sup> CFU mL <sup>-1</sup>	11 CFU mL <sup>-1</sup>	Selective, Recoveries85.7% 105.5%	Milk	[298]	

Green and red QD Carbon particle	A p	<i>V. parahaemolyticus</i> & <i>S. Typhimurium</i>	FL/FRE T	50-10 <sup>6</sup> CFU m L <sup>-1</sup>	6 <i>V. parahaemolyticus</i> 35 <i>S. Typhimurium</i> CFU m L <sup>-1</sup>	Selective, 35 <i>S. Typhimurium</i> CFU m L <sup>-1</sup>	Shrimp Chicken breast	
CdSe/ZnS QD	MNP 20-150 nm CdSe/ZnS	A p	<i>avian influenza virus</i> (AIV) H5N1	FL QCM	2 <sup>-12</sup> -2 <sup>6</sup> HAU 20 μL <sup>-1</sup>	0.4 HAU	Specific, Rapid, 96.6% Signal after 3 month	[299]
AgNCs	AgNCs synthesized reducing AgNO <sub>3</sub> with NaBH <sub>4</sub> stabilized by the DNA template spherical in shape with the average size of about 2 nm	A p	<i>S. Typhimurium</i>	FL	10 <sup>5</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	50 CFU mL <sup>-1</sup>	Specific, detecting viable S. from heat-denatured, RSD 3% n=5	Chicken meat [300]
Carbon nanotube	Gel green fluorescence ssDNA aptasensor based on carbon nanotubes for detection of anthrax protective antigen	A p	recombinant protective antigen domain 4 (rPAD4) of <i>Bacillus anthracis</i>	FL	5-80×10 <sup>-6</sup> g L <sup>-1</sup>	20 ×10 <sup>-6</sup> g L <sup>-1</sup> 62.5 ×10 <sup>-6</sup> g L <sup>-1</sup> purified and unpurified rPAD4 protein		Mouse serum sample [301]
AuMNPs CdTe QD	AuNPs coated Maghemite nanoparticles maghemite 10 nm	D N A Pr ob e		FL	5×10 <sup>-8</sup> 2×10 <sup>-4</sup> mol L <sup>-1</sup>	10 <sup>-10</sup> mol L <sup>-1</sup>	Selective, immobilization of particles with bound targets using a force of the external magnetic field	Serum [302]
MNPs& AuNPs	MNPs close to complete super paramagnetic: AuNPs citrate 15 nm		DNA probes <i>Shigella spp</i>	FL	10 <sup>1</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	2.1 ng mL <sup>-1</sup> or 90 CFU mL <sup>-1</sup>	Selective	[303]
CdTe quantum dots (QDs)	Mono disparity & spherical morphology particle size of about 3.5 nm		DNA probe Human Papillomavirus 18 virus (HPV18) gene	FL/FRE T	1.0×10 <sup>-9</sup> 5×10 <sup>-8</sup> mol L <sup>-1</sup>	2×10 <sup>-10</sup> mol L <sup>-1</sup>	Selective	DNA from positive patient [304]
CdTe QDs	Red and green CdTe QDs were 8 ± 3 nm and 5± 2 QDs showed its fluorescence emission quantum yields up to 25%		antisense-E6-HPV16 (HPV-probe) Gag-HIV (HIV-probe) E6-HPV16 &Gag-HIV.	FL	2×10 <sup>-10</sup> mol L <sup>-1</sup>	-10 <sup>-7</sup> 5×10 <sup>-11</sup> mol L <sup>-1</sup>	Simultaneous detection of HPV & HIV, Selective, The results were approved by PCR and electrochemistry	DNA sample of Patient [305]
lysozyme-AuNCs	Microwave-assisted synthesis AuNCs 3-6nm green and facile one-pot microwave-assisted method red fluorescence gold nanocluster	lysozyme- <i>E. coli</i>		FL	2.4×10 <sup>4</sup> 6.0×10 <sup>6</sup> CFU mL <sup>-1</sup>	2.0×10 <sup>4</sup> CFU mL <sup>-1</sup>	Green and facile one-pot microwave-assisted method for the synthesis of red fluorescence Au nanoclusters	[306]
Guanidine @UCNP	GDN@UCNPs have good environment adaptability UCNPs hexagonal size of around 20 nm.	-	<i>Shigella flexneri</i> / <i>Vibrio parahaemolyticus</i> / <i>Staphylococcus aureus</i> / <i>Listeria monocytogenes</i>	FL	10 <sup>3</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	1.3×10 <sup>2</sup> CFU m L <sup>-1</sup>	Facile, Low cost, Without the need for biomolecule	Tap water, Beef, Milk [307]
Au nanocluster	AuNCs25 gold atom 2.29 nm	-	<i>E. coli</i>	FL	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	89 CFU mL <sup>-1</sup>	Selective, Fast	[308]
Au nanocluster	BSA-AuNCs, IHS-AuNCs & Lys-AuNCs	-	<i>Pseudomonas aeruginosa</i> , <i>Bacillus natto</i> , <i>E. coli</i> , & <i>Acetobacter aceti</i>	FL	-	-	5 different bacteria and 9 protein identified by the sensor array	- [309]
Carbon dots	CD modified with vancomycin, CD synthesized with citric acid and urea and microwave treatment, spherical and well-dispersed 2.27 ± 0.3 nm	-	<i>Gram positive</i> <i>L. S. aureus</i>	FL	3.18×10 <sup>2</sup> - 1.59×10 <sup>8</sup> CFU mL <sup>-1</sup>	9.40×10 <sup>4</sup> CFU mL <sup>-1</sup>	Low cost and easy to perform	Orange juice [310]
Carbon dots	Green synthesis of QDs from papaya peel Water-soluble 2- 6 nm, and ethanol 8 -18 nm soluble CD with 18.98 and 18.39 %Quantum yield respectively, Water-soluble CDs better imaging compared ethanol because of smaller size and more quick entrance to cell	-	<i>E. coli</i>	FL	-	9.5×10 <sup>4</sup> CFU mL <sup>-1</sup>	Low cost and easy to perform Water CDs and ethanol-CDs are of low toxicity to HeLa cells	- [311]
Carbon dots	CDs@ colistin well dispersed in water 2-5 nm the quantum yield of 7.56%.	-	<i>E. coli</i>	FL	3.18×10 <sup>2</sup> - 2.44×10 <sup>8</sup> CFU mL <sup>-1</sup>	578 CFU mL <sup>-1</sup>	Simple, cheap, Selective	Urine, Apple juice, Tap water [312]

collect the solution containing both nanoparticle and poly (ethylene glycol) in a pipette tip respectively. Poly (ethylene glycol) medium caused separation Pt/MNC- *E. coli O157: H7* nanoparticle from Free Pt/ MNC nanoparticle. 10 CFU mL<sup>-1</sup> *E. coli O157: H7* could detect by color change (oxidation TMB with nanoparticle) [136]. More colorimetric biosensors are presented in Table 2. Lateral flow and the immunochromatographic assays will discuss in a separate part in section 3.6.

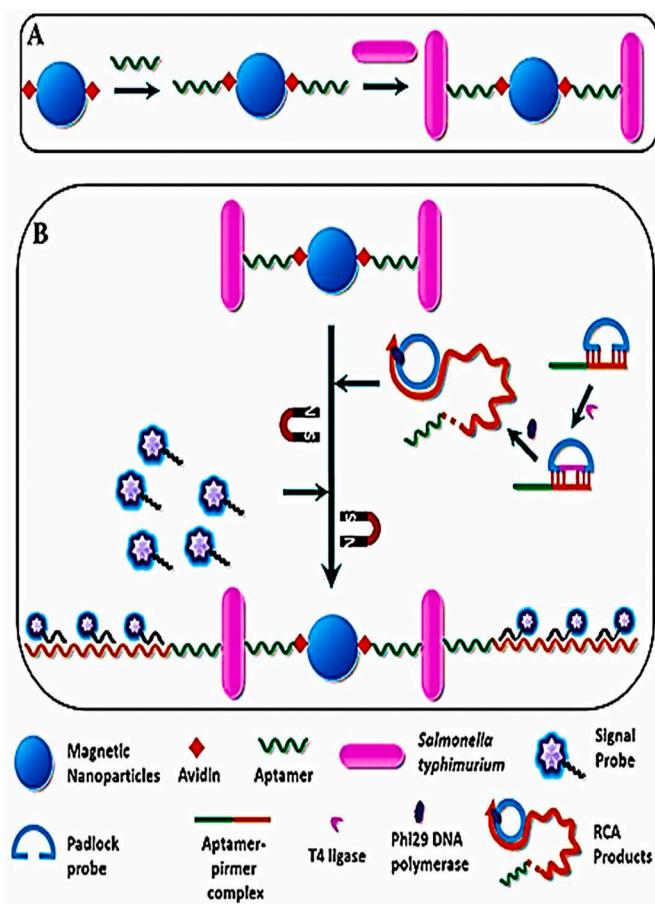
### 3.3. Fluorescence biosensors

Fluorescence-based sensors have attracted great interest in presenting high sensitivity, low detection limit, rapid response, low signal-to-

noise ratio, simple instrumentation, and cost-effectiveness. A variety of fluorescence nano-materials such as metal nanoparticles, quantum dots (QDs), carbon dots (CDs), graphene oxide, dye-doped silica nanobeads, nanotubes, and upconversion nanoparticle have been explored in sensing application [137]. Signal generation in fluorescence sensing has been carried out following a variety of strategies including fluorescence quenching (“turn-off”), fluorescence enhancement (“turn-on”), ratio-metric(calculating intensity ratio of two or more fluorescence signal), fluorescence resonance energy transfer (FRET), photoinduced electron transfer (PET) and metal enhanced fluorescence(MEF) [138].

Specific separation and detection of *E. coli O157: H7* was reported with the use of a magnetic immunoassay and QDs labeling. The capture

of the bacteria was performed in a double layer quartz channel using the immune magnetic nanoparticle trapped on the channel wall by a high gradient magnetic field. Following labeling with QD, the formed sandwich complex was collected from the channel and the fluorescence intensity of the sample measured. This was shown to relate linearly with log concentration of bacteria in the range  $8.9 \times 10^0$ – $8.9 \times 10^5$  CFU mL $^{-1}$  [139]. The schematic of this procedure is illustrated in Fig. 4A. Aptamer coated magnetic particles labeled with QD modified complementary DNA were used in a competitive assay for the detection of *S. Typhimurium*. In the presence of the target bacteria, the aptamer-DNA label duplex was cleaved due to the aptamer-target interaction resulting in free CdTe QD labeled DNA in solution. After magnetic separation, the QDs signal in solution was recorded and demonstrated to be linearly related to  $10$ – $10^{10}$  CFU mL $^{-1}$  of log pathogen concentration [140]. Application of QD and upconversion nanoparticle (UCNP) modified, respectively, with *S. Typhimurium* and *S. aureus* aptamers allowed the simultaneous detection of these two food pathogens. After attachment of the two luminescent nanoparticles to the magnetic bead contains a partially complementary DNA sequence, the fluorescence emission spectra obtained at 500 nm for QD and 800 nm for UCNPs was recorded; the addition of the target pathogens to the solution containing the sensing complex resulted in the loss of fluorescence labels with a subsequent decrease in the overall signal intensity of the magnetic beads (recorded after magnetic separation). The reduced signal correlated to the log concentration of the bacteria in the  $50$ – $10^6$  CFU mL $^{-1}$  interval [141]. Liu and coworkers extracted *V. parahaemolyticus*-specific egg yolk antibody and used these to decorate AuNPs. Interaction of CdSe/ZnS QD which was decorated with 3-mercaptopropionic acid (carrying carboxylic acid group) and AuNPs, modulated by COOH groups in the QDs and Au $^0$  atom in AgNPs lead to fluorescence quenching of QD. A small amount of free AuNPs was also added that enable a higher rate of quenching. In the presence of *V. parahaemolyticus* due to the interaction of bacteria and antibody, IgY-AuNPs aggregated freeing in this way the QDs and increasing the fluorescence response of them. The system was applied to the sensitive detection *V. parahaemolyticus* in food samples [142]. Composite of AuNPs, gallic acid-iron oxide nanoparticle, and graphene, modified with hemagglutinin (HA) antibody, was employed as an integrated magnetic and plasmonic surface for the capturing and detection of the influenza virus. The addition of QDs, also decorated with hemagglutinin (HA) antibody, caused the formation of a sandwich complex and the fluorescence of QD increased linearly with virus concentration. The detection limits of  $7.27 \times 10^{-12}$  g L $^{-1}$  and  $6.07 \times 10^{-9}$  g L $^{-1}$  were obtained in deionized water and Human serum respectively [143]. Quaternary alloyed CdZnSeTe QDs which can emit in the NIR range is an attractive option for the construction of low detection limit biosensor due to the low photonic-absorption of biological molecule in this area. 2 copies mL $^{-1}$  of Influenza virus H1N1 RNA were detected with molecular beacons (MB) modified with the aforementioned QDs by fluorescence enhancement signal transduction mechanism. In the absence of the target sequence, QDs and MB in close proximity resulted in an efficient quenching of their IR signal. Hybridization of target viral RNA with the loop sequence of the MB probe formed DNA/RNA heteroduplex which enhanced the fluorescence of QD by inducing the separation of the two particles [144]. The schematic of this procedure is illustrated in Fig. 4B. A FRET aptasensor consisting of AuNPs modified with aptamer (acceptor) and UCNPs conjugated with complementary DNA (donor) was designed for the ultrasensitive detection of *Escherichia coli* ATCC 8739, in food sample and water, down to 5 CFU mL $^{-1}$ . Attachment of bacteria to aptamer led to the separation of the UCNPs-CDNA from the AuNPs- Aptamer resulting in the loss of the FRET effect with subsequent recovery of the fluorescence signal of the donor [145]. Hollow silica nanospheres (HSNs) encapsulated with fluorescein and covered with polymer layers of poly (acrylic acid) and poly (dimethyldiallyl ammonium chloride) were synthesized and used in the development of a label in a sandwich magnetic immunoassay, following modification with Ab $_2$  antibodies, for the detection of *E. coli*.



**Fig. 5.** The pattern of chemiluminescence biosensor A) Aptamer attachment and bacteria capturing B) Rolling circle amplification and Co $^{2+}$ -enhanced signal probes with permission ref [153].

Fe $_{3}$ O $_{4}$ @SiO $_{2}$ @PAA-Ab $_1$  were used for the capture of the bacteria and the separation of the formed sandwich. Signal was generated, following the collection of the formed sandwich, by dissolution, with sodium hydroxide, of the HSNs label, and release of the internal fluorescein. The fluorescence intensity of the released fluorescein is related to the *E. coli* O157: H7 concentration down to 4 CFU mL $^{-1}$  [146]. FRET aptasensor based on AuNCs-Vancomycin and aptamer modified AuNPs as the energy donor and acceptor was developed for the sensitive detection of *S. aureus*. In the presence of *S. aureus* fluorescence intensity change ( $\Delta F = F_0 - F$ ) increased linearly with concentration  $20$ – $10^8$  of *S. aureus* [147]. Two main bacteria responsible for sepsis (*S. aureus*, *E. coli*) were separated and detected in mouse rat blood. In the presented work, Fe $_{3}$ O $_{4}$  nanoparticles modified with chlorin e6 and *S. aureus* and *E. coli* aptamers were designed for the early and sensitive ( $10$  CFU mL $^{-1}$ ) detection of the two pathogens by fluorescence microscope. Moreover, chlorin e6, photosensitizers, applied and extracorporeal blood disinfection in mouse rats' blood was achieved with photodynamic therapy [148]. The schematic of this procedure is illustrated in Fig. 4C. A fluorometric aptasensor was fabricated by employing aptamer modified carbon nanoparticle and GO as fluorescent and quencher respectively for the sensitive detection of *P. aeruginosa*. In the presence of bacteria, the fluorescence signal was increased due to binding aptamer and bacteria that caused distance between the aptamer-carbon nanoparticle and Go, the recorded signal related linearly to  $10^1$ – $10^7$  CFU mL $^{-1}$  concentration of bacteria with a LOD of 9 CFU mL $^{-1}$  [149]. An Aptasensor consisting of MNP-Aptamer and UCNPs-CDNA was developed to detect, low concentration ( $58$  CFU mL $^{-1}$ ), *E. coli* in pork meat sample. MNP-Aptamer and UCNPs-CDNA mixed and incubated for short time. Then, the addition of

**Table 4**

Chemiluminescence and photoluminescence biosensors, Abbreviation chemiluminescence (CL) [313–315].

Nanomateri al	Nanomaterial remark	Bi o	Pathogen	Method	Linear range	LOD	Remark	Real sampl e	Ref
GO oxide/Fe <sub>3</sub> O <sub>4</sub>	Fe <sub>3</sub> O <sub>4</sub> Co-precipitation	A p	<i>E. coli</i> O157:H7	CL	10 <sup>4</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup> .	4.5×10 <sup>3</sup> CFU mL <sup>-1</sup> .	Easy-to-use, Fast	Skim milk	[313]
Co <sup>2+</sup> /N-(aminobutyl)-N-(ethylisoluminol) functional flowerlike gold Fe <sub>3</sub> O <sub>4</sub> -NH <sub>2</sub> MNP Fe <sub>3</sub> O <sub>4</sub> -NH <sub>2</sub> MNP 50nm AuNPs nm carrier for CL reagents & the catalyst to enhance CL intensity Good dispersibility and morphology with an		A p	<i>S. Typhimurium</i>	CL	32–3.2×10 <sup>6</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective	Pork sampl es	[151]
Co <sup>2+</sup> /N-(aminobutyl)-N-(ethylisoluminol) functional flowerlike gold Fe <sub>3</sub> O <sub>4</sub> -NH <sub>2</sub> MNP, WS <sub>2</sub> ; Nanosheet Co <sup>2+</sup> /N-(aminobutyl)-N-(ethylisoluminol) functional flowerlike gold as a donor & WS <sub>2</sub> Nanosheet acceptor		A p	<i>Staphylococcus</i> <i>aureus</i>	CL	50–1.5×10 <sup>5</sup> CFU mL <sup>-1</sup>	15 CFU mL <sup>-1</sup>	Recoveries of CL ranged from 94.7 to 102.0% 3.72% n=7	Pork paste	[153]
AuNPs	AuNPs <sub>c</sub> 20nm, Biotin, and Luminol attached to the surface of AuNPs	A b	<i>Hepatitis B</i> surface antigen	CL	1.7×10 <sup>2</sup> – 1.92×10 <sup>6</sup> g L <sup>-1</sup>	3×10 <sup>10</sup> g L <sup>-1</sup>	Specific, Sensitive	Human Serum	[314]
AuCNTs, Cystamine coated CdTe QD	AuNPs 20nm	A b	Influenza viruses H <sub>3</sub> N <sub>2</sub>	photolu minesce nce	50–10 <sup>5</sup> PFU mL <sup>-1</sup>	50 PFU mL <sup>-1</sup> . 0.1 pg mL <sup>-1</sup>	Selective, Sensitive	Clinic al sampl e	[315]

*E. coli* release UNCP- CDNA particle due to the specific interaction of aptamer and target. Following the separation of *E. coli*-MNP-aptamer complex by the magnet, the fluorescence signal of the solution was recorded which was increased by increasing concentration of bacteria [150]. More fluorescence biosensors are demonstrated in table 3

### 3.4. Chemiluminechence and photoluminescence biosensors

Chemiluminescence (CL) takes advantage of the conversion of chemical energy into light. As a result, this method does not require sample radiation reducing in this way interfering phenomena such as light scattering, source instability, providing in this way high signal to noise ratio and high sensitivity; moreover, the readout instruments are less complicated than other optical systems [151,152].

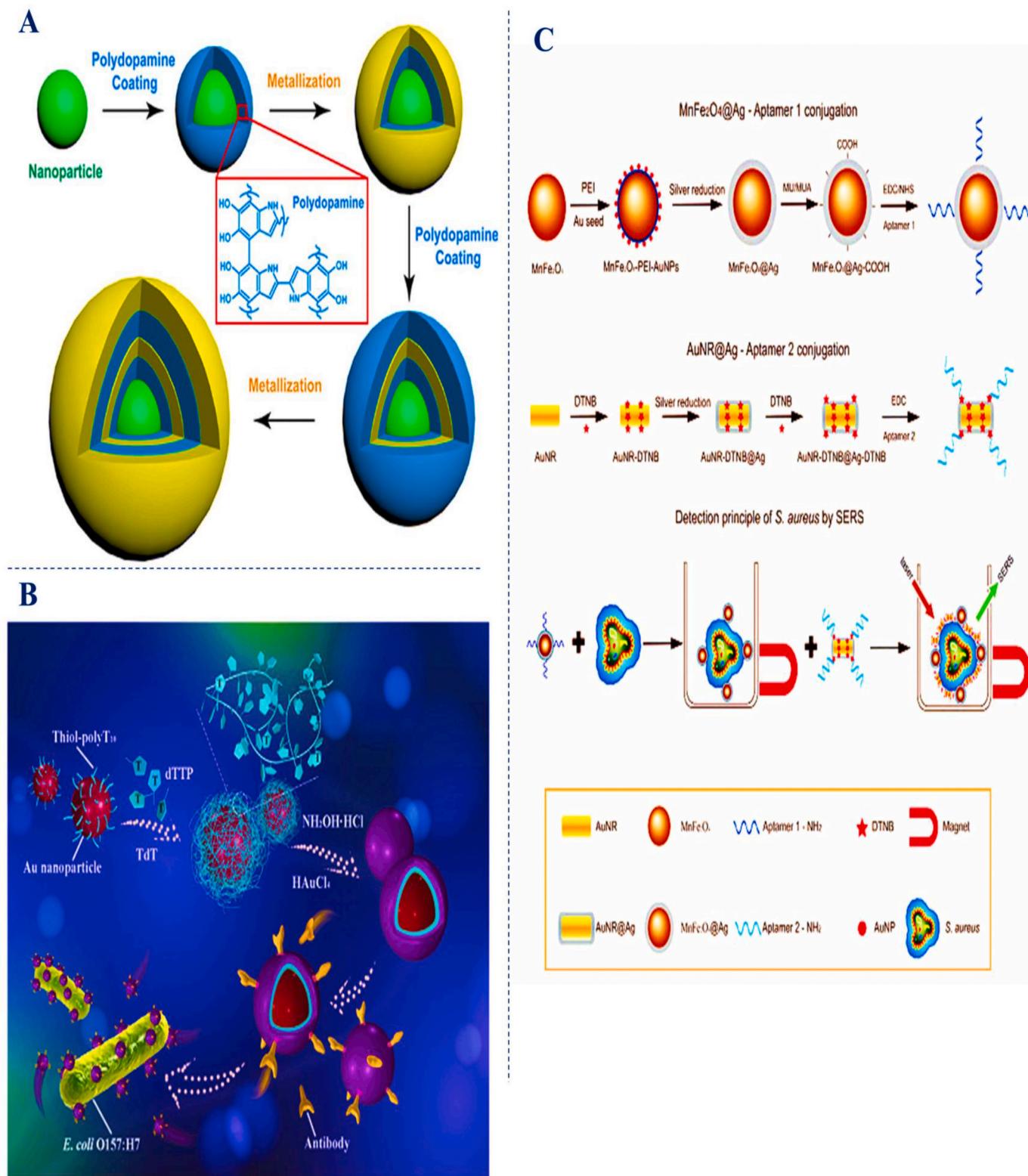
Hao et al. reported on the steady-state Chemiluminescence apta-sensor, based on the rolling circle amplification (RCA) method, for the sensitive detection of *S. Typhimurium*. In the proposed strategy target bacteria are captured using MNP immobilized aptamer; captured *S. Typhimurium* can then interact with the RCA product to form a sandwich complex. Finally, the formed sandwich was labeled using a Co<sup>2+</sup> enhanced N- (aminobutyl)-N-(ethylisoluminol) (ABEI) functional flowerlike gold nanoparticles (AuNFs)-cDNA. Chemiluminescence signal was then generated by the addition of P-Indophenol and H<sub>2</sub>O<sub>2</sub> applied to form ABEI- AuNFs-PIP-H<sub>2</sub>O<sub>2</sub> Chemiluminescence system. The proposed sensor was shown to respond linearly in the concentration range between 3.2 and 3.2 × 10<sup>6</sup> CFU mL<sup>-1</sup> [153]. The schematic of the presented assay was shown in Fig. 5. More examples of Chemiluminechence and photoluminescence methods are presented in Table 4.

### 3.5. Surface-enhanced Raman scattering (SERS) biosensors

Surface-enhanced Raman scattering is a powerful sensing tool; this presents several advantages including high sensitivity, reduced analysis time, portability as well as the ability for multiplex detection. SERS technology takes advantage of the enhancement of the Raman spectroscopy signal when performed at the surface of noble metal nanomaterials; signal enhancements of several orders of magnitude (10<sup>7</sup>–10<sup>14</sup>) were reported using this method. SERS is a fascinating method for the identification (fingerprinting) and detection of microorganisms because of its non-destructive nature. Nanomaterials like noble metal colloids, nanospheres, core-shell, gold-coated magnetic nanoparticle, nano aggregates, and bimetallic nanomaterial have been reported as promising materials for SERS detections using both labels (indirect) and label-free (direct) approaches. Labels (SERS reporters) are specific

organic molecules like, Rhodamine B, 4-Nitrothiophenol 4-mercaptopurine (4-MPY), 4-amino thiophenol (4-ATP), 4-mercaptopbenzoic acid and 5,5-dithiobis-2-nitrobenzoic acid (DTNB) that are characterized by well-known Raman signals. Plasmonic nanoparticles decorated with Raman reporter molecules and biorecognition elements have been used to design SERS biosensors [154,155].

In a study reported by Wang et al. [156], positively charged polyethyleneimine was self-assembled on Fe<sub>2</sub>O<sub>4</sub> magnetic nanoparticle; the surface of the MNP was modified, taking advantage of electrostatic interaction, with Au seeds that were then chemically grown to generate rough core/shell (Au/MNPs) label. To enable the selective capture of *S. aureus*, Au/MNPs were modified by the *S. aureus* antibody. On the other side, Au nanorods -DTNB modified with antibodies were used to complete the sandwich assay. SERS intensity at Raman peak of 1331 cm<sup>-1</sup> showed a linear relationship with the logarithmic concentration of the bacteria in the range of 10<sup>1</sup>–10<sup>5</sup> cells mL<sup>-1</sup> [156]. Concanavalin A (Con A) is a lectin that presents the ability to specifically interact with the terminal α-D-mannosyl and α-D-glucosyl groups present on the surface of all bacteria. Kearns et al. reported on a SERS assay based on the use of Ag-MNPs core-shell system modified with Con A. The reported core-shell system enabled the capture and collection of bacteria from samples. With the help of three different SERS reporters, the authors demonstrated the multiplex detection of *S. Typhimurium*, methicillin-resistant *S. aureus*, and *E. coli* down to 10 CFU mL<sup>-1</sup> [157]. Zhou et al. carried out extensive studies on the design and fabrication of multifunctional nano-gapped nanoparticles (NNPs). In their works polydopamine (PDA) coating was used to deposit, in a controlled way (e.g. using PDA of different thicknesses), Au shells on a variety of core nanomaterials as spherical AuNPs, anisotropic Au nanorods (AuNRs), metal-organic frameworks (MOFs), and magnetic polymer nanoparticles (MAgNPs). The produced nanogap structures were reported as a carrier for Rhodamine-B or 4-nitrophenol Raman tags in bacteria sandwich assays; magnetic nanoparticle coated with PDA and 2 layers of 15 nm Au nanoshell and modified with 4- nitrophenol (Raman tag) enabled the sensitive detection of *E. coli* O157: H7 down to concentration 100 CFU mL<sup>-1</sup>. In the same work, the authors showed that the Raman signal could be significantly enhanced (ca. 5 folds) by the increase in the number of nanoshell layers (2 vs 1). The schematic of preparation nanogap structures is illustrated in Fig. 6A [158]. Thiol-poly adenine or Thiol-poly thymine was immobilized on the AuNps surface. Then, terminal deoxynucleotidyl transferase which was mediated incorporation of the chain of identical nucleotides utilized to synthesize nucleotide with different lengths due to change of elongation time (0 min –16 h). The prepared nanoconjugate worked as a nano seed to synthesis a gold



**Fig. 6.** A) Schematic of the synthesis of nanogapped nanoparticles based on the polydopamine coating, B) Schematic of the synthetic sequence in producing hT-DENPs for recognition of *E. coli* O157: H7 C) Synthesis of silver-coated magnetic nanoparticles and their conjugation with aptamer 1, synthesis of core–shell plasmonic nanoparticles (AuNR–DTNB @Ag – DTNB) and their conjugation with aptamer 2 Operating principle for *S. aureus* detection.

layer to immobilize the antibody on it. Among different nanogap structures, the optimum one (16 h) with poly adenine was selected for the rest of the investigations. The so prepared particles were used as a Raman reporter in a bacteria immune assay that was shown to enable the detection of 2 CFU mL<sup>-1</sup> of *E. coli* [159]. The schematic of the

proposed immunosensor was illustrated in Fig. 6B. AuNPs decorated with 4, 4'-dipyridyl, and coated with silica layers were shown to allow bacterial detection in complex systems being stable for up to 50 h and not being prone to aggregation. The SERS immunosensor based on such nanoparticles was shown to detect 10 CFU mL<sup>-1</sup> of *E. coli* [160].

**Table 5**

Surface-enhanced Raman scattering (SERS) biosensors, Abbreviation Loop-mediated isothermal amplification (LAMP), Membrane filter-assisted (MFA), Recombinase polymerase amplification (RPA) [316–340].

Nanoparticle	Nanoparticle performance	Biomolecule	Pathogen	method	Linear range	Detection limit	Remark	Real sample	Ref
AuNPs	distance between cy5 Raman tag less than 2 nm,	highly serovar - specific DNA region within the <i>Salmonella</i> gene <i>Salmonella Enterica</i>	SERS/ LAMP	6.6×10 <sup>0</sup> -6.6 ×10 <sup>6</sup> CFU m L <sup>-1</sup>	66 CFU m L <sup>-1</sup>	Great selectivity		Milk	[316]
AuMNPs, Au nanorods	AuMNPs highly uniform in size and shape with good magnetic responsiveness, stability, and high SERS activity, Superparamagnetic properties.	Ab <i>S. aureus</i>	SERS	10 <sup>1</sup> -6.6 ×10 <sup>4</sup> cell mL <sup>-1</sup>	10 cell mL <sup>-1</sup>	Capture and fast separation of bacteria, Sonochemically assisted hydroxylamine seeding growth method for the facile synthesis of AuMNPs for the first time.			[156]
Functionalized polymeric MNPs, AuNPs	FPMNPs aqueous dispersion and storage stability up to one year at room temperature 140nm, AuNPs 31nm	Ab <i>S. Typhimurium</i>	SERS	10 <sup>1</sup> -10 <sup>7</sup> cell mL <sup>-1</sup>	10 cells mL <sup>-1</sup>	Specific Good reproducibility	Cottage cheese Egg whit Juice		[317]
MNP@Ag@ConA	MNP CO-precipitation method, Ag shell with the glucose reduction method	Ab <i>S.Typhimurium</i> , <i>methicillin-resistant S. aureus</i> <i>E.coli</i>	SERS	10 <sup>2</sup> -10 <sup>4</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Easy to operate,3 different Raman reporter for simultaneous detection of 3 bacteria	-		[157]
MNP, AuNPs /SWCNT	MNP CO-precipitation method, 10-20nm	Ab <i>E.coli</i>	SERS	10 <sup>2</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, AuNP@SWCNT photothermal killing effect	Water		[318]
Au shell	AuNPs <sub>c</sub> covered by poly adenine or thymine followed by Terminal deoxynucleotidyl transferase addition and Au Layer	Ab <i>E. coli O157: H7</i>	SERS	10 <sup>1</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	2 CFU mL <sup>-1</sup>	Specific	Apple juice		[159]
Multi shell plasmonic nanogap nanoparticle, AuNRs, MNPs	AuNRs seed-mediated method Magnetic particle co-precipitation method AuNPs <sub>c</sub> covered with polydopamine and Au shell, AuNPs covered with polydopamine and Rhodamine B	Ab <i>E.coli O157:H7</i>	SERS	10 <sup>2</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Selective, Adopted nanogap size by the thickness of Polydopamine, the photothermal killing effect	River water		[158]
AuNPs silica-coated	AuNPs 30nm silica shell 4 nm Magnetic beads 100nm	Ab <i>E.coli O157: H7</i>	SERS	10-10 <sup>8</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Specific, AuNPs silica-coated stabilized signal 90 days	River water		[160]
AuMNPs, Au@MBA@Van	Au citrate + MBA SERS tag +Vancomycin 52± 3 nm, Fe <sub>3</sub> O <sub>4</sub> @Au MNPs synthesized by a PEI-mediated seed growth method 200 ± 6 nm, Au shell 25 ± 5 nm 90-day stability, (MS)Fe <sub>3</sub> O <sub>4</sub> @Au-aptamer MNPs are	Ap <i>S. aureus</i>	SERS	10-10 <sup>7</sup> cell mL <sup>-1</sup>	3cell mL <sup>-1</sup>	Selective, Vancomycin low price.	Milk, orange juice human blood		[163]

decreased bout 8.8% after aptamer coating									
Spiny Gold Nps	The Spiny Gold Nps absorption peak red-shifted to 850 nm and the particle size was 200~120 nm	Ap <i>S. Typhimurium</i>	SERS	10 <sup>1</sup> -10 <sup>5</sup> CFU m L <sup>-1</sup>	4 CFU m L <sup>-1</sup>	Excellent sensitivity, Selective Recoveries 96.55% -105.45%	Pork		[164]
SiO <sub>2</sub> @ Au core-shell	Silica Stöber method 200 ±19 nm	Ap <i>Vibrio parahaemolyticus</i>	SERS	10 <sup>1</sup> -10 <sup>6</sup> CFU m L <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Specific, cyanine dye Raman tag	Shrimp		[319]
GO oxide wrapped Fe <sub>3</sub> O <sub>4</sub> @Au	GO oxide wrapped Fe <sub>3</sub> O <sub>4</sub> @Au as separation source and SERS substrate AuNPs <sub>c</sub> 13 nm Fe <sub>3</sub> O <sub>4</sub>	Ap <i>Vibrio parahaemolyticus</i>	SERS	1.4 ×10 <sup>2</sup> -1.4 ×10 <sup>6</sup> CFU m L <sup>-1</sup>	14 CFU mL <sup>-1</sup>	Selective	Salmon samples		[320]
AuNPs <sub>c</sub>	AuNPs <sub>c</sub> 40 nm Capture DNA	Ap <i>E.coli O157: H7</i>	SERS	10 <sup>2</sup> -10 <sup>6</sup> CFU m L <sup>-1</sup>	10 <sup>1</sup> ground beef 10 <sup>2</sup> CFU mL <sup>-1</sup>	Specific,10 <sup>2</sup> CFU mL <sup>-1</sup> ground beef	Ground beef		[321]
AuNPs		Ap <i>S. enteritidis</i>	SERS	1.4 ×10 <sup>2</sup> -1.4 ×10 <sup>7</sup> CFU m L <sup>-1</sup>	10 <sup>3</sup> CFU m L <sup>-1</sup>	Specific, Fast (3h)	Fresh spinach		[322]
MNPs, AuNPs	MNPs Co-precipitation method AuNPs <sub>c</sub> 15nm	Ap <i>S. Typhimurium</i> & <i>S. aureus</i>	SERS	10 <sup>2</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	35& 15 CFU mL <sup>-1</sup>	<i>S. aureus</i> & <i>S. Typhimurium</i> Selective	Commercial pork		[323]
AgNPs	AgNPs NaBH <sub>4</sub>	Ap <i>S.aureus</i>	SERS	10 <sup>1</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	1.5 CFU mL <sup>-1</sup>	Absence of Raman marker			[161]
MnFe <sub>2</sub> O <sub>4</sub> , AgMNPs, Au Nanorod	MnFe <sub>2</sub> O <sub>4</sub> Superparamagnetic 170nm binary solvent Solvothermal AgMNPs 200nm DTNB-labeled inside-and-outside plasmonic NPs (DioPNPs) that exhibits high SERS conjugates with target recognition molecules (Ab or Ap)	Ap <i>S. aureus</i>	SERS	10 <sup>1</sup> -10 <sup>5</sup> cells mL <sup>-1</sup>	10 cells mL <sup>-1</sup>	Specific, Raman signal of DioPNPs is 10 times			[162]
Ag@Au nanoshells	AuNPs + AgNO <sub>3</sub> + Ascorbic acid AuNPs 15 nm+ Ag shell 4nm	Ap <i>S. Typhimurium</i>	SERS	15-1.5×10 <sup>6</sup> CFU mL <sup>-1</sup>	15 CFU mL <sup>-1</sup>	Selective	Milk		[165]

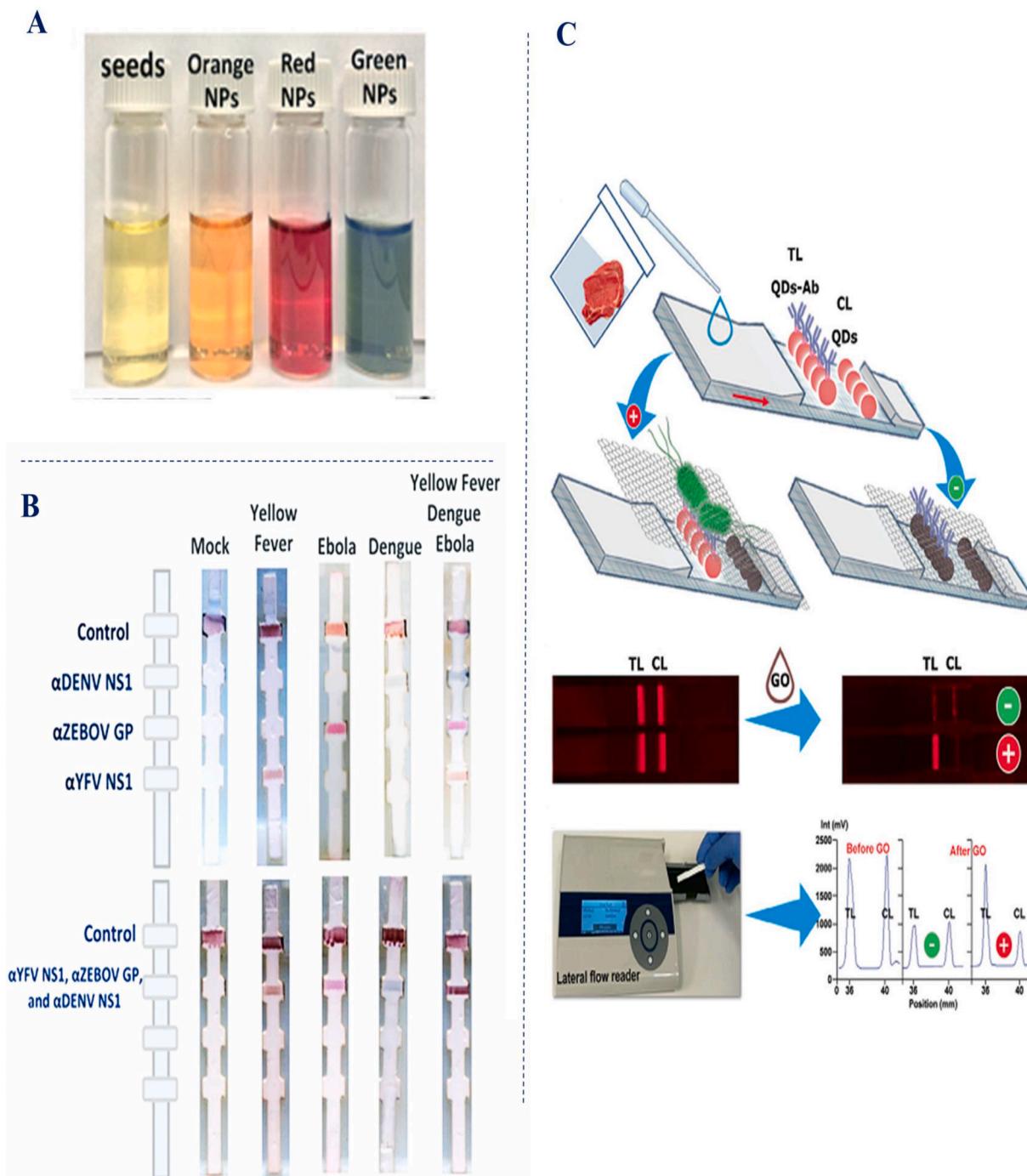
MNP, AuNPs	Magnetic nanoparticles 15–20 nm co-precipitation method, AuNPs <sub>C</sub> MNP increased the Raman intensity due to thin Au shell surfaces, enabled the MGPNPs to maintain their magnetic property but also obtain a much better dispersibility.	Ap	<i>S. Typhimurium</i>	SERS	10 <sup>1</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	5 CFU mL <sup>-1</sup>	Selective	Pork	[324]
Ag NRs	Ag NRs synthesized in a custom-built electron beam evaporator using the oblique angle deposition technique	Ap	<i>S. Typhimurium</i>	SERS	-	10 <sup>8</sup> CFU mL <sup>-1</sup>	Label-free	-	[325]
MNP, AuNPs	MNP 30 nm	Ab	<i>E. coli O157:H7</i>	SERS/ MFA	10–500 CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective	Ground beef	[326]
Fe <sub>3</sub> O <sub>4</sub> @Au nanoparticle, Fe <sub>3</sub> O <sub>4</sub> @AuV an	Fe <sub>3</sub> O <sub>4</sub> @Au nanoparticle uniform in size and morphology, with 250 nm, 25 nm Au shell Fe <sub>3</sub> O <sub>4</sub> @AuVan , Vancomycin layer 1.5nm	Ap	<i>E. coli &amp; S. aureus</i>	SERS	50–10 <sup>5</sup> <i>E. coli</i> 20–10 <sup>5</sup> <i>S. aureus</i> cell mL <sup>-1</sup>	20 and 50 cells mL <sup>-1</sup> for <i>S. aureus</i> & <i>E. Coli</i>	Selective, Rapid capture of bacteria, Fe <sub>3</sub> O <sub>4</sub> @Au effectively capture bacteria in comparisonFe <sub>3</sub> O <sub>4</sub> @Au	Urine	[327]
AuNPs, AuNds	AuNds 35 nm DNA carrier, AuNPs 15 nm asymmetric gold nanodimers uniform & monodisperse stabilized SERS activity	Ap	<i>S. Typhimurium</i>	SERS	10 <sup>2</sup> –10 <sup>7</sup> CFU mL <sup>-1</sup>	35 CFU mL <sup>-1</sup>	Selective,	Milk	[328]
Silica Encapsulated Nanotags, MNP	Raman reporter +Au layer+ Silica shell 1015nm, SERS nanotags with shielding shells to avoid the leaching of Raman reporters and capture biomolecules as well as to	Ab	<i>West Nile virus (WNV)</i> envelope (E) protein, <i>Rift Valley fever virus (RVFV)</i> nucleocapsid (N) protein, Capsular antigen fraction I (FI) <i>Yersinia pestis</i>	SERS	10 <sup>3</sup> –10 <sup>5</sup> g L <sup>-1</sup>	10 <sup>11</sup> g mL <sup>-1</sup> in 20% bovine serum	The thickness of the silica shell is important, Simultaneous detection	Fetal bovine serum	[329]
MNPs/SiO <sub>2</sub> AuNPs	Large AuNPs <sub>C</sub> 51 nm+ MBA MNPs CO-participation	DN A sequence tags	bacterial DNA	SERS/ PCR	10 <sup>-20</sup> –10 <sup>12</sup> mole	zeptomole (4.1×10 <sup>-11</sup> ) g/L	Excellent sensitivity	-	[330]
Fe <sub>3</sub> O <sub>4</sub> @Ag-Van MNPs, Au@Ag core-shell	Superparamagnetic Fe <sub>3</sub> O <sub>4</sub> particles ~200 nm vancomycin-modified Ag capture agent and SERS substrate, 30 s separation Au@Ag core-shell NPs 60 nm	<i>E. coli BL21</i> , <i>S. aureus</i> , &MRS aureus		SERS	5×10 <sup>2</sup> Cell m <sup>-1</sup>	5×10 <sup>2</sup> Cell mL <sup>-1</sup>	11-mercaptopentadecanoic acid Raman Tag bifunctional linker thiol and carboxylic acid	Milk Blood	[331]
AgNPs	Microwave heating method, 50–70 nm AgNPs greatly enhanced the SERS signal of bacteria	- <i>E. coli O157: H7</i> , <i>S. aureus</i> <i>Salmonella</i>		SERS	-	10 <sup>7</sup> CFU mL <sup>-1</sup>	Without the need for biomolecule	-	[332]
AgNPs	AgNPs Hydroxylamine in basic solution	<i>E. coli</i>		SERS	1×10 <sup>2</sup> –1×10 <sup>7</sup> Cell mL <sup>-1</sup>	10 <sup>5</sup> Cell m L <sup>-1</sup>	Without the need for biomolecule	-	[333]
Fe <sub>3</sub> O <sub>4</sub> @PEI @Au, Fe <sub>3</sub> O <sub>4</sub> @Au@PEI, Au@Ag NPs	Fe <sub>3</sub> O <sub>4</sub> 300nm solvothermal, Au shell 3 nm super-paramagnetic, Au@AgNPs 60 nm better SERS activity. The Au@Ag-bacteria complexes showed an improvement about 1.5 and 3.2 times stronger than 60 nm Au NPs and 50 nm Au NPs	<i>E. coli &amp; S. aureus</i>			10 <sup>3</sup> –10 <sup>7</sup> cells mL <sup>-1</sup>		Rapid, Simple operation	Tap water, Milk	[334]
AgNPs		<i>S. enterica</i> and <i>Listeria monocytogenes</i>			3.5×10 <sup>4</sup> 3.5×10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	3-mercaptophenylboronic acid capture and reporter for bacteria		[335]
Ag collides	Synthesis of AgNPs within a bacterial culture with NaBH <sub>4</sub>	<i>P. aeruginosa</i> , <i>L. mono cytogenes</i> & <i>L. innocua</i> and <i>MRSA-86</i> , <i>MRSA-35</i>			<i>L. innocua</i> 10 <sup>1</sup> –10 <sup>3</sup> CFU mL <sup>-1</sup>		Pre-treatment of cells with Triton X-100 considerably enhanced the sensitivity		[336]
AgNPs	In situ synthesis of Ag nanoparticles with close contact with the bacterial membrane.	<i>E. coli</i> , <i>M. morganii</i> , <i>E. lactis</i> , <i>L. casei</i>		SERS			Rapid, Low cost, single-cell detection		[337]
AgNPs, AuNPs	AgNPs <sub>C</sub>	<i>E.coli</i> , <i>Shewanella putrefaciens</i> <i>Pseudomonas aeruginosa</i>		SERS			Discriminate between the three yeast species		[338]
AgNPs	AgNPs <sub>C</sub> 100nm	<i>Yeast</i> , <i>Saccharomyces cerevisiae</i> , <i>Dekkera bruxellensis</i> <i>Wickerhamomyces anomalus</i>		SERS			Discriminate between the three yeast species	Beer	[339]
Au@Ag	Au@Ag self-assembled mussel shell AgNPs 40 ± 12.5 nm with 5nm shell	<i>E. coli</i> , <i>S.aureus</i> , & <i>P. aeruginosa</i>		SERS			Discriminate between the three bacteria		[340]

Label-free detection of *S. aureus* was performed by the use of specific aptamers and by in situ synthesis, using the aptamer as a template, on AgNPs. The SERS spectra of *S. aureus*-aptamer/AgNPs were shown to be significantly higher than those recorded in the absence of the aptamer. The system was reported to have a low LOD of 1.5 CFU mL<sup>-1</sup> [161]. Ag coated magnetic nanoparticle (AgMNPs) and AuNR –DTNB@Ag-DTNB (DioPNP) as SERS tags were reported in single-cell detection of *S. aureus*. These were modified with aptamer 1 and 2, respectively, to form a dual sandwich structure; the Raman signal of DioPNPs was shown to be 10 times stronger than those of the AuNR –DTNB due to double-layer DTNB [162]. The schematic of preparation nanoparticle and bacteria detection was shown in Fig. 6C. Dual enhancement SERS aptasensor was developed by Pang et al. and applied to the detection of *S. aureus*; Fe<sub>3</sub>O<sub>4</sub>@Au –Aptamer nanoparticles were used for the specific recognition/capture of the target bacteria with SERS signal, due to the captured target, generated at the Au layer. The second element of the assay was AuNPs modified with-4 mercaptobenzoic acid (Raman label) and vancomycin (for bacteria capturing). The proposed biosensor was shown to allow detection of the pathogen in a wide range of concentrations (10–10<sup>7</sup> CFU mL<sup>-1</sup>) with a LOD of 3CFU mL<sup>-1</sup> [163]. Ma et al. developed a

sensitive aptasensor with the aid of pointy-like AuNPs that have unique field enhancement especially at the tip of the branches. The pointy-like AuNPs were used, following functionalization with 4- mercaptobenzoic acid and thiolated aptamer, as SERS label in ELISA plate-based aptamer assay. ELISA plates were modified with biotinylated aptamer used to capture *S. Typhimurium*. The proposed detection approach was shown to detect down to 10 CFU mL<sup>-1</sup> of *S. Typhimurium* [164]. Duan et al. reported on an aptasensor for the detection of *S. Typhimurium*; this was based on the use of an Au@Ag core-shell nanoparticle modified with aptamer 1 as a signal enhancer and of aptamer2-X-rhodamine as SERS label. The fabricated biosensor enables sensitive detection of *S. Typhimurium* down to 15 CFU mL<sup>-1</sup> [165]. More example of SERS bio/sensor are summarized in Table 5.

### 3.6. Lateral flow-based biosensors

Lateral-flow immunoassay (LFIA) also known as lateral-flow immunochromatographic (LF- ICA) tests are simple, rapid, and portable systems which enable qualitative, non-invasive point of care detection (POC) with wider applications in medical diagnostic [166].



**Fig. 7.** A) AgNPs for multiplexed detection, B) Multiplex detection of dengue, yellow fever, and Ebola viruses with permission ref [173] C) Fluorescent lateral flow immunoassay ref [177]. (For interpretation of the references to color in this figure legend, the reader is referred to the Web version of this article.)

Conventional LFA system suffers from low sensitivity; nevertheless, in recent times several approaches for the improvement of LFA have reported including gold enhancement, enzyme labeling, increased loading of antibody, and use of various nanomaterials like QD, UCNPs, peroxidase-like nanoparticle, and graphene oxide. Besides coupling LFIA systems with instrumental reading (fluorescence, colorimetric, SERS, and magnetic focus) enabled to obtain more reliable and quantitative data [167,168].

An innovative approach, combining gold-coated  $\text{Fe}_3\text{O}_4$  magnetic nanoparticle (Au/MNCs) and lateral flow filters, was shown to allow the detection of  $10^3 \text{ CFU mL}^{-1}$  *Salmonella* from milk samples. In the proposed assay *Salmonella* was captured in milk samples using antibody

fragment modified Au/MNCs; the formed complex following separation was resuspended in a buffer solution. The assay was then performed by immersing one end of the lateral stripe in the solution; if on one end the free Au/MNCs were able to move in the LFI strip and accumulating in the test line, the *Salmonella*-Au/MNCs were remained in the solution because of their big size; pressing of the nitrocellulose membrane was key factor in the success of the proposed assay. The colour of the test line decreased with increasing pathogen concentration because of a decrease in free Au/MNCs nanoparticle [169]. The same author also developed a similar strategy for the visual detection of *E. coli* O157: H7 ( $10^3 \text{ CFU mL}^{-1}$ ) [170]. Cho et al. developed the lateral-flow immunochromatographic LF-ICA system based on AuNP-biotin and streptavidin -HRP

complex. A mixture of tetramethylbenzidine and H<sub>2</sub>O<sub>2</sub>, applied in the cross-flow direction of the antigen-antibodies complex, was used as a reagent for the generation of the colorimetric signal; the proposed assay allowed the naked eyes detection of *Cronobacter sakazakii* down to 10<sup>2</sup> CFU mL<sup>-1</sup> [171]. The size and uniformity in the shape of AuNPs were shown to influence the sensitivity of LFIA. Cui and co-workers investigated the effect of size and uniformity of AuNPs and pre-incubation of AuNPs/antibody/*E. coli O157: H7* on the visual and instrumental reading of a colorimetric test strip. The authors synthesized AuNPs with size 20, 28, 35, 43, 54 nm; AuNPs of 35 nm showed the best results. If on one end smaller nanoparticles, due to their smaller steric hindrance, interacted more effectively with bacteria, larger ones offered better optical signal, and the 35 nm nanoparticles had plasmonic resonance wavelength 525 which was the same as the wavelength light of light source in the reader. Nanoparticles with lower polydispersity indices (PDI) provide a superior limit of detection [172]. Yen et al. synthesized AgNPs with different size and colour modified with *dengue* virus (DENV) NS1 protein, *Yellow Fever Virus* (YFV) NS1 protein, and *Zaire strain Ebola* virus glycoprotein (ZEBOV) for single and multiplex detection of these viruses. The visual detection of the three viruses demonstrates to be possible down to 150 ng mL<sup>-1</sup> [173]. Different AgNPs with the prepared stripe is depicted in Fig. 7A. Jin et al. investigated 15 different AuNPs decorated polystyrene microparticle (Au-Ps) with different sizes (0.1–0.3, 0.46) μm and Au coverage (5, 10, 25, 50, 75%) to find the nanoparticle with the highest sensitivity for visual detection of *E. coli O157: H7*. The optical density of the different nanoparticles was used as the physical parameter to discriminate between the manufactured particles and to select the best one. According to the optical density values, the UV-vis spectrums, and the TEM images, (50Au-0.46 PS) was expected to provide the better result nevertheless average grayscale analysis by image J software showed that lower LOD could be achieved with the (10Au-0.46 PS) nanoparticle. The same authors also studied the effect of signal enhancement by reduction of HAuCl<sub>4</sub> with hydroxylamine; this enhancement step lowered the LOD down to 100 CFU mL<sup>-1</sup> and average 1.44 folds improvement for each concentration [174]. Eu (III) doped polystyrene nanoparticles were demonstrated in immunochromatographic assay for detection of *E. coli O157: H7*. EuNP-6 carbon chain (CC), EuNP-200CC, EuNP-1000CC and EuNP-streptavidin were compared; EuNP-Streptavidin, due to their better ability to load antibodies, showed better sensitivity and wider linear range [175]. Ren et al. reported on the combination of Fe<sub>3</sub>O<sub>4</sub>/Au core-shell and HRP signal enhancement for the detection of pathogens. The use of the described system, in combination with an external magnet to slow down the movement of the labeled target which enabled the interaction of pathogens and antibody more readily, allowed detection limits of 25CFU mL<sup>-1</sup> *E. coli O157: H7* or *S. Typhimurium* [176]. A fluorescence lateral flow immunosensor was designed by taking advantage of the quenching properties of GO (acceptor) and the fluorescence ability of QDs (donor). In the proposed assay the test line contained CdSe@ZnS QDs/anti-*E. coli O157:H7* antibody while the control line only had bare QDs. In the absence of target bacteria, addition of GO resulted in the quenching of the fluorescence of the QDs in both the test and the control lines. By contrast, capturing of *E. coli O157: H7* with antibody, increase the distance between QD and GO, therefore reducing the ability of GO to quench the QDs; the intensity of the residual fluorescence was shown to be proportional to the pathogen concentration in the range 50–10<sup>5</sup> CFU mL<sup>-1</sup> with LOD 57 CFU mL<sup>-1</sup> [177]. The schematic of the assay was showed in Fig. 7B. Surface positive nitrogen-rich carbon nanoparticle (pNC), prepared from urea by calcination and etching reaction, were shown to have a good ability to adsorb/capturing bacteria. In the proposed LFIA monoclonal antibodies present on the test line were used to capture the complex made between the pNC and the bacteria; the intensity of back color of the test line was shown to relate to the concentration of *S. enteritidis* in the 10<sup>2</sup>–10<sup>8</sup> CFU mL<sup>-1</sup> interval [178]. Plasmonic LFIA with improved performances based on the AuNPs aggregation induced by polyethyleneimine (BPEI)

released from liposome was reported by Ren et al. for the sensitive detection of *E. coli O157: H7*. Targeted bacteria were incubated with AuNPs modified with an antibody; following a certain time BPEI-loaded liposomes were added to the solution. The obtained mixture was added to the strip followed by the addition of free-AuNPs. The added nanoparticles were then captured at the test line by the BPEI present on the surface of the bacteria/AuNPs complex resulting in the generation of a colorimetric signal due to the aggregation effect. The proposed immunoassay could detect 100 CFU mL<sup>-1</sup> of *E. coli O157: H7* [179].

The peroxidase activities of some nanomaterials lead to select them as a prominent candidate for use in LFIA with improved sensitivity. For example, the peroxidases activity of Pt-AuNPs modified with antibody were explored to generate, upon reaction with TMB, a colored band in correspondence of the test line. The proposed assay enabled the detection of 10<sup>2</sup> CFU mL<sup>-1</sup> *E. coli O157: H7* [180]. Mesoporous core-shell Pd@Pt with the help of TMB and H<sub>2</sub>O<sub>2</sub> was used for the simultaneous detection of *S. enteritidis* and *E. coli O157: H7* with LFIA and smartphone detection down to 20 and 34 CFU mL<sup>-1</sup> of *S. enteritidis* and *E. coli O157: H7* while the detection limit of the system for both pathogens was 10<sup>6</sup> CFU mL<sup>-1</sup> in the absence of TMB and H<sub>2</sub>O<sub>2</sub> system. AuNR@Pt with peroxidase activity toward TMB and H<sub>2</sub>O<sub>2</sub> and also SERS properties with 4-mercaptopbenzoic acid Raman reporter were explored for dual recognition, LFIA and SERS methods; the two methods allowed, respectively, the detection of *C. jejuni* in the 10<sup>2</sup>–10<sup>6</sup> and 10<sup>2</sup>–5 × 10<sup>6</sup> CFU mL<sup>-1</sup> concentration ranges [181]. Use of Fe<sub>3</sub>O<sub>4</sub> MNP as a nano-enzyme probe was reported by Han et al. for the LFIA detection of the glycoprotein of the *Ebola* virus. The ability of the MNPs to catalyze the conversion of 3, 3'-Diaminobenzidine (DAB) in the presence of H<sub>2</sub>O<sub>2</sub> allowed the detection of the glycoprotein at 1 ng mL<sup>-1</sup> concentration which is 100 times lower than that achievable with the standard strip method (1 ng mL<sup>-1</sup>) [182]. Bimetallic Pd-Pt nanoparticles were also explored as nano-enzyme for the conversion of TMB and H<sub>2</sub>O<sub>2</sub> in LFIA for the detection of *E. coli O157: H7*; the proposed assay showed LOD of 10<sup>2</sup> CFU mL<sup>-1</sup> [183]. Fe<sub>3</sub>O<sub>4</sub>/Au-polyethyleneimine nanoparticles-casein-antibody complex were applied to capture *E. coli*, followed by cleavage of casein by rennet enzyme to release *E. coli*. Released bacteria were loaded on strip modified with DTNB labeled AuNPs and AuNPs modified with the antibody on the conjugate pad and test line respectively. The intensity of the test line was recorded which was increased with concentration 10<sup>1</sup>–10<sup>7</sup> CFU mL<sup>-1</sup> *E. coli* with LOD 0.52 CFU mL<sup>-1</sup> [184]. Luo et al. compared 4 different nanoparticles including AuNPs, QD, Fluorescent nanoparticles (FNP), and Eu(III) chelate nanoparticles for application in the immunochromatographic assay for the detection of *E. coli O157: H7*. The EuNP showed the highest fluorescence intensity, larger Stokes shift, and sharpest emission profile, therefore the EuNP showed the highest sensitivity and fluorescent nanoparticles and EuNP provided the wider linear range. It needs to be noted that fluorescent nanoparticles were shown to require less antibody per strip [185]. Aptamer mediated strand displacement approach was reported for the sensitive detection of *E. coli O157: H7*. In the reported assay a reporting aptamer and a capture probe, modified with biotin, were used. Target bacteria were incubated with the two aptamers and extracted using streptavidin magnetic beads. After the separation of the magnetic complex, the reporting aptamer was amplified with the isothermal strand displacement amplification method and detected with LFIA. This method enabled visual detection of 10 CFU mL<sup>-1</sup> *E. coli O157: H7* [186]. Mesoporous silica nanoparticles, embedded with TMB and covered with amin-aptamer gate sequence, were employed for the sensitive detection of *L. monocytogenes*. The addition of bacteria caused the release of TBM due to their specific interaction with the aptamers. Released TMB interact with HRP enzymes trapped in the test line to produce the visible blue color. The aptasensor could obtain LOD of 53 cells in 1 mL sample [187]. AuNPs modified with 4-mercaptopbenzoic acid (as a Raman tag), Ag layer, streptavidin and BSA were used as a label for the detection of *L. monocytogenes* and *S. enterica* DNA sequences. In the proposed assay the targeted DNA sequences were amplified, before their LFIA detection, using recombinase polymerase amplification method. The proposed assay allowed the visual and SERS detection of the targeted sequences with LOD of 27 and 19 CFU mL<sup>-1</sup>. Visual qualitative detection was made possible via the aggregation of the

**Table 6**

Lateral flow-based biosensor detection Lateral flow immunoassay (LFIA) immunochromatographic immunoassay (ICA) Fluorescence Lateral flow immunoassay (FLFIA), Immunomagneticseparation (IMS), Polymerase chain reaction (PCR), Strand displacement amplification (SDA) [341–374].

Nanomaterial	Nanomaterial remark	B i o	Metho d	Linear range	LOD	Remark	Real sample	Ref
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 30nm	A <i>Pseudomonas syringae</i> pv. b <i>maculicola</i>	LFIA	10 <sup>5</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>5</sup> CFU mL <sup>-1</sup>	Broccoli and radish seeds, the IC strips sensitivity of 5 × 10 <sup>5</sup> & 10 <sup>6</sup> CFU mL <sup>-1</sup>	Broccoli & Radish seeds	[341]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> average diameter 35 nm	A <i>Cronobacter sakazakii</i> b	LFIA	10 <sup>7</sup> -10 <sup>3</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Specific, Enhancement 100 times in LOD and 3h in enrichment time by increasing the amount of Ab by conjugating with concentrated AuNPs in the test line	Powdered infant formula	[167]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 24nm in diameter	A <i>E.coli O157:H7</i> b	LFIA	1.65 × 10 <sup>4</sup> - 3.3 × 10 <sup>6</sup> CFU mL <sup>-1</sup>	1.58 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	Selective, Simultaneous detection of <i>E.coli O157:H7&amp; Aflatoxin M1</i>	Milk	[342]
AuNPs <sub>C</sub>	15nm	A <i>Listeria</i> b	LFIA	10 <sup>7</sup> -10 <sup>3</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Selective, Sensitivity enhance 1order of magnitude by strip reader 1-9 CFU mL <sup>-1</sup> after 8h	Milk	[343]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 15 nm	A <i>Listeria monocytogenes</i> b	LFIA	-	3.7 × 10 <sup>6</sup> CFU mL <sup>-1</sup>	Specific, 8- 12 h enrichment for Milk sample	Milk	[344]
Au/MNCs	MNC synthesized with the hydrothermal method, MNCs and Au/MNCs 160 and 180 nm, AuNPs <sub>C</sub> 14nm	A <i>E.coli O157:H7</i> b	Size-sorting LFIA	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Selective, Novel strategy by squeezing a nitrocellulose membrane to reduce its thickness	Milk	[170]
Au/MNCs	MNC synthesized with the hydrothermal method, MNCs and Au/MNCs 160 and 180 nm, AuNPs <sub>C</sub> 14nm	A <i>Salmonella</i> b	Size-sorting LFIA	10 <sup>3</sup> -10 <sup>5</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Selective, Novel strategy by squeezing a nitrocellulose membrane to reduce its thickness	Milk	[169]
AuNPs	AuNPs Size 40 nm	A <i>E.coli O157:H7</i> b	LF- ICA	10 <sup>3</sup> -10 <sup>6</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> Visual CFU mL <sup>-1</sup>	Selective, Applying AuNP-Avidin and biotin-HRP complex. HRP catalyzes TMB oxidation	Ground beef	[171]
AuNPs	AuNPs <sub>C</sub> solutions with different PDI and size 20,28,35,43,54	A <i>E.coli O157:H7</i> b	LFIA	10 <sup>3</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>3</sup> CFU mL <sup>-1</sup>	Uniform size AuNPs size and separated incubation of AuNPs/antibody/ <i>E. coli</i> O157:H7 complex to improve sensitivity 35nm best 10 <sup>7</sup> CFU mL <sup>-1</sup> Specific reader10 <sup>7</sup> CFU mL <sup>-1</sup>	-	[172]
MNPs	MNPs hydrothermal method	A glycoprotein (GP) of the <i>Ebola Virus</i> b	LFIA	10 <sup>-6</sup> -10 <sup>-3</sup> g L <sup>-1</sup>	10 <sup>-6</sup> g L <sup>-1</sup> 240 PFU mL <sup>-1</sup> pseudo-	Selective,100 fold increase in sensitivity with help MNPs intrinsic peroxidase-like activity feature & DAB	Clinical sample	[182]
EBOV								
Au magnetic bifunctional nanobeads	30 nm	A <i>Salmonella choleraesuis</i> b	ICT	1×10 <sup>8</sup> 5×10 <sup>5</sup> CFU mL <sup>-1</sup>	5×10 <sup>5</sup> CFU mL <sup>-1</sup>	Selective	Milk	[345]
AuNPs	AuNPs <sub>C</sub> 20 nm	A <i>Salmonella</i> b	LFIA	10 <sup>4</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>4</sup> CFU mL <sup>-1</sup>	Selective	River water	[346]
AuNPs <sub>C</sub>	Fe <sub>3</sub> O <sub>4</sub> AuNPs <sub>C</sub> 30 nm	A <i>Enterobacter cloacae</i> b	IMS/ ICT	10 <sup>3</sup> -10 <sup>7</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	With the use of Immunomagnetic beads, the target bacteria were separated and enriched, Selective,10 times higher than that obtained by direct detection with ICTS	Water	[347]
AuNPs	AuNPs 15 nm	A <i>Shigella boydii</i> & ( <i>E. coli</i> O157:H7)	ICA	10 <sup>6</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>6</sup> CFU mL <sup>-1</sup>	With pre-incubation 4CFU mL <sup>-1</sup>	Bread Milk Jelly	[348]
AuNPs	A 28 nm and 45 nm with spherical shape were demonstrated to be the optimal dual pair	A <i>E.coli O157: H7</i> b	LFIA	10 <sup>5</sup> -10 <sup>3</sup> CFU mL <sup>-1</sup>	1.14×10 <sup>3</sup> CFU mL <sup>-1</sup>	Comparing 4 nanoparticles with size 18,28,45,80 as dual pair.	Reduced-fat milk	[349]
AuNPs, QD, FNP, EuNPs	QD,FNP, EuNps size 21, 11, 182, 155 nm EuNP had the highest fluorescence intensity	A <i>coli O157: H7</i> b	LF- ICA	1.0 × 10 <sup>-5</sup> - 1.0 × 10 <sup>6</sup> 2.5 × 10 <sup>3</sup> - 1.0 × 10 <sup>6</sup> 2.5 × 10 <sup>2</sup> - 2.5 × 10 <sup>5</sup> 2.5 × 10 <sup>2</sup> - 2.5 × 10 <sup>5</sup> For GNP-ICA, QD-ICA, FNP-ICA, GNP-ICA, QD- ICA, FNP-ICA EuNP-ICA	2.5 × 10 <sup>4</sup> 5 × 10 <sup>3</sup> 1.0 × 10 <sup>3</sup> 5.0 × 10 <sup>2</sup> CFU mL <sup>-1</sup> For GNP-ICA, QD-ICA, FNP-ICA, GNP-ICA, QD- ICA, FNP-ICA EuNP-ICA	Specific, Comparing 4 nanoparticle The EuNP-ICA showed the highest sensitivity, FNP-ICA need the least Ab per strip, The FNP-ICA, and EuNP-IC broader linear range FNP-ICA and EuNP-ICA better resistance in the milk sample	Milk	[185]
Au-decorated polystyrene particles (Au-PS)	AuNPsC Mixing AuNPs With PS and boiling them to 100 °C AuNPs 20 nm diameter size dispersed on PS particle surface	A <i>O157: H7</i> b	LFIA	5×10 <sup>2</sup> -1×10 <sup>6</sup> CFU·mL <sup>-1</sup>	500 CFU·mL <sup>-1</sup>	Selective, lowered LOD 1×10 <sup>2</sup> CFU mL <sup>-1</sup> with signal amplification with w 1% HAuCl <sub>4</sub> and 10mMNH <sub>4</sub> OH-HCl 1:10, Au-PS particles with 0.46 μm PS with 50% Au coverage has maximum OD but the one with 10% coverage has better LOD	Ground beef	[174]

AgNPs	Three AgNPs by seed-mediated growth method, Size: orange 30 ± 7 nm, Red 41 ± 6 nm, green= 47± 8 nm	A <i>Dengue, Yellow Fever protein, and Ebola viruses</i> b <i>Listeria monocytogenes</i>	LFIA g L <sup>-1</sup>	150-500 ×10 <sup>-6</sup> 150 ×10 <sup>-6</sup> g L <sup>-1</sup>	Multiplex detection with three different AgNPs, Selective	Human serum	[173]
Carboxyl functionalized superparamagnetic beads	140 nm	A <i>Listeria monocytogenes</i> b	LFIA/ Magnetic assay reader	10 <sup>4</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup> 10 <sup>4</sup> CFU mL <sup>-1</sup>	Specific	Food samples	[350]
Core-shell red silica nanoparticle	Monodispersed SiO <sub>2</sub> covered reactive red 126 NPs were prepared by Stöber methods spherical morphology and good dispersity diameter 216 nm	A <i>E.coli O157: H7</i> b	LFIA CFU mL <sup>-1</sup>	4.5 ×10 <sup>4</sup> -4.5 ×10 <sup>5</sup> CFU mL <sup>-1</sup>	Specific 4.5 ×10 <sup>6</sup> -CFU mL <sup>-1</sup>	Milk and pork sample	Pork and milk sample [351]
Nanobeads	Magnetic nanobeads	A <i>E.coli O157: H7</i> b	FLFIA	Selective, Stroger signal with magnetic nanobeads Detect 1,5, 25, and 125 CFU of in 25 mL milk in 8, 7, 6, and 5 h.		Milk	[352]
Eu (III)-doped polystyrene nanoparticle	EuNP & EuNP-linker size 158 , 159 , 158 , 161,159 nm,	A <i>E. O157: H7</i> b	FICA	8×10 <sup>1</sup> - 1.59×10 <sup>5</sup> CFU·mL <sup>-1</sup>	Selective, carboxyl-modified EuNPs with 4 linkers. EuNP-SA-ICA best result	Milk	[175]
EuNPs	EuNPs 103 nm ± 1.4 nm, EuNP-McAb 114 nm ± 1.7 nm	A <i>E. coli O157:H7</i> b	FLFIA	10 <sup>4</sup> -10 <sup>7</sup>	Strip& quantitative detection was 10 <sup>4</sup> & 5 × 10 <sup>2</sup> CFU mL <sup>-1</sup> with the test-strip reader	Specific, Pre-incubation for 8 h in meat and 6 h Milk LOD1CFU mL <sup>-1</sup>	Beef, Pork, Chicken [353]
Fe <sub>3</sub> O <sub>4</sub> /Au core-shell AuNPs.	Au core-shell nano-35-50 nm AuNP40 nm.	A <i>E. coli O157:H7</i> , b <i>S.Typhimurium</i>	LFIA/ Magnetic focus	0-10 <sup>2</sup> CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup>	Specific	Pineapple juice [176]
CdSe/ZnS-COOH	Water-soluble QD Spherical 3nm	A <i>E. coli O157: H7</i> b	FLFIA	10 <sup>4</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	10 <sup>9</sup> CFU mL <sup>-1</sup>	Specific, 35-day stability, fast detection time 12 min	Milk [354]
GO & r GO	GO, rGO and conjugated rGO sheets 1 μm,rGO produce higher color intensity	A <i>E. coli O157: H7</i> b	LFIA	10 <sup>7</sup> -10 <sup>9</sup> CFU mL <sup>-1</sup>	10 <sup>5</sup> CFU mL <sup>-1</sup>	Specific, rGO10 <sup>6</sup> Drinking water 10 <sup>7</sup> Milk CFU mL <sup>-1</sup>	Milk Drinking water [355]
Go, CdSe@ZnS	CdSe@ZnS QDs, of size14 ± 2 nm &	A <i>E. coli O157: H7</i> b	FLFIA	50 to 10 <sup>5</sup> CFU mL <sup>-1</sup>	57 CFU mL <sup>-1</sup>	Selective,178 and 133 CFU g <sup>-1</sup> or mL <sup>-1</sup> real sample, The single Water	Minced beef Water [177]
maximum emission wavelength at ≈665 nm							
GO, CdSe@ZnS	Optimal QDs concentration be 1.5 nM The optimum GO concentration was determined to be 90 μg mL	A <i>E. coli</i> b	FLFIA	10 <sup>10</sup> <sup>5</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Antibody approach reduced the assay cost to 60% less, 10 CFU mL <sup>-1</sup> in standard buffer and 100 CFU mL <sup>-1</sup> in bottled water and milk	Bottled Milk water [356]
MoS <sub>2</sub> & GO	MoS <sub>2</sub> & Go nanosheets well defined laminar morphology with a uniform size of around 200 and 500 nm	A <i>Salmonella enteritidis</i> b	LFIA/I CT	MoS <sub>2</sub> 10 <sup>3</sup> -10 <sup>8</sup> GO 10 <sup>4</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	MoS <sub>2</sub> 10 <sup>3</sup> GO 10 <sup>4</sup> CFU mL <sup>-1</sup>	Specific	Drinking water Watermelon juice [357]
CdSe/ZnSQDs /Fe <sub>3</sub> O <sub>4</sub>	carboxyl-poly styrene/acrylamide copolymer nanospheres 250 nm coated with Fe <sub>3</sub> O <sub>4</sub> and QDs repeatedly and with an outer silica layer	A <i>S.Typhimurium</i> b	LFIA/ Magnetic reader	1.88×10 <sup>4</sup> - 1.88 × 10 <sup>7</sup> Typhi, 3.75×10 <sup>3</sup> FL	1.88 × 10 <sup>4</sup> CFU mL <sup>-1</sup>	Selective, Simple a magnetic assay reader, range of 1.88	Tap water Milk, Fetal bovine serum Whole blood [358]
Streptavidin AuNPs	40nm	A <i>Salmonella</i> and <i>E.coli</i> b Amplicon	Quadruple taggin g PCR/N AFA	<i>E.coli</i> 7×10 <sup>8</sup> g L <sup>-1</sup> <i>Salmonella</i> 9.5×10 <sup>5</sup> g L <sup>-1</sup>	-	Simultaneous detection, Selective 6.6×10 <sup>7</sup> g L <sup>-1</sup> g L <sup>-1</sup> for <i>E.coli</i> ( in presence of high concentrations of <i>Salmonella</i> ), 1.17×10 <sup>6</sup> L <sup>-1</sup> For salmonella (in presence of High concentration of E.coli	[359]
Surface positively charged nitrogen-rich Carbon nanoparticles(g-C <sub>3</sub> N <sub>4</sub> )	g-C <sub>3</sub> N <sub>4</sub> pNC generate a signal and capture Bacteria	A <i>Salmonella enteritidis</i> b	LFIA	10 <sup>2</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>2</sup> CFU mL <sup>-1</sup>	Good sensitivity, Broad linear range, Specific, Single Ab, Recoveries ranging from 85% - 110%.	Potable water, Coleslaw cabbage Salad, watermelon juice Purple cabbage [178]
NaYF <sub>4</sub> :Yb <sup>3+</sup> Er <sup>3+</sup>	50 nm	A <i>E. coil</i> O157: H7 S. <i>paratyphi A</i> , <i>S. paratyphi B</i> , <i>S. paratyphi C</i> , <i>S. enteritidis</i> , <i>S. Typhi</i> , <i>S. choleraesuis</i> , <i>V. cholera O1</i> ,	LFIA	10 <sup>4</sup> -10 <sup>8</sup> CFU mL <sup>-1</sup>	10 <sup>7</sup> or 10 <sup>8</sup> CFU mL <sup>-1</sup>	Simultaneous detection of 10 pathogens	Dairy products Marine products Beverages Snacks Meats [360]

<i>V. cholera O139, V. parahaemolyticus</i>								
Pd-Pt nanoparticles	Pd-Pt nanoparticles size 40 nm	A <i>E. coli O157: H7</i> b	LFIA	$10^3\text{-}10^6$ CFU mL <sup>-1</sup> $10^3\text{-}10^6$ CFU mL <sup>-1</sup> $9 \times 10^2$ CFU mL <sup>-1</sup> in Milk	$1 \times 10^2$ CFU mL <sup>-1</sup>	Selective, Amplifying signal With peroxidase-like catalytic activities of Pd-Pt NP& TMB & H <sub>2</sub> O <sub>2</sub>	Milk	[183]
Au-Pt bimetallic	Porous 50 nm.	A <i>E. coli O157: H7</i> b	ICA	$10^2\text{-}10^8$ CFU mL <sup>-1</sup>	$10^2$ CFU mL <sup>-1</sup>	Amplifying signal With - peroxidase-like catalytic activities of Au-Pt NP& TMB & H <sub>2</sub> O <sub>2</sub>	-	[180]
Pd@Pt	Pd@Pt mesoporous core-shell 135 nm	A <i>S. enteritidis</i> & <i>E. coli O157:H7</i>	LFIA/ Smart phone	$10^2$ CFU mL <sup>-1</sup>	Selective, Simultaneous, 30 day stability, $10^6$ CFU mL <sup>-1</sup> without TMB & H <sub>2</sub> O <sub>2</sub>	30 day without Milk Ice cream	[361]	
MNPs, AuNPs	MNPs 180 nm AuNPs 13 nm	Ab DNA probe <i>Listeria monocytogenes</i>	IMS/L FIA	$10^3\text{-}10^7$ CFU mL <sup>-1</sup> $3.5 \times 10^3$ CFU mL <sup>-1</sup> $3.5 \times 10^4$ CFU g <sup>-1</sup> Lettuce	$10^3$ CFU mL <sup>-1</sup>	Selective, Eliminate dead cell, Less amount of Ab, based on biotin exposure strategy	lettuce samples	[362]
Magnetic beads	1000, 500, 200 , 130 nm size	A <i>E.coli</i> b	Visual/ CO/ LFIA	$10^3\text{-}10^7$ CFU mL <sup>-1</sup>	$10^5$ CFU mL <sup>-1</sup>	Selective, 6-month stability	River water	[363]
AuNR@Pt	AuNR@Pt length & diameter of around 65 and 15 nm, with Pt layer with the spiky shape	A <i>Campylobacter jejuni</i> b	SERS/ LFIA	$10^2\text{-}10^6$ CFU mL <sup>-1</sup> and $10^2$ $-5 \times 10^6$ CFU mL <sup>-1</sup>	$75$ CFU mL <sup>-1</sup> and $50$ CFU mL <sup>-1</sup>	Selective, Easy operation, Rapid	Milk Chicken	[181]
Casein modified Fe <sub>3</sub> O <sub>4</sub> /Au-PEI	AuNPs <sub>c</sub> 2-6 nm	A <i>E. coli</i> b	SERS/ LFIA	$10^2\text{-}10^7$ CFU mL <sup>-1</sup>	LOD 0.52 and LOQ = 1.57 CFU mL <sup>-1</sup>	Selective, Wide linear range	Urine Milk	[184]
AuNPs	AuNPs 25nm	A <i>S.Typhimurium</i> b	ICA/E LISA	$1.3 \times 10^5$ $2 \times 10^6$ CFU mL <sup>-1</sup>	$1.25 \times 10^5$ Milk $1.25 \times 10^5$ CFU mL <sup>-1</sup>	Specific, Comparison 10 different antibody	Milk	[364]
carboxylate-modified CdSe/ZnS core@shell, Fe <sub>3</sub> O <sub>4</sub> spherical 100 nm, Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> 250 nm,Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> @ NH <sub>2</sub> , Fe <sub>3</sub> O <sub>4</sub> @SiO <sub>2</sub> @QD Stöber255 nm	QD 10 nm, Fe <sub>3</sub> O <sub>4</sub> solvothermal method	A <i>E. coli O157: H7</i> b	FLFIA/ IMS	$2.5 \times 10^2$ CFU mL <sup>-1</sup> Visual $5 \times 10^5$ CFU mL <sup>-1</sup> Visual	$2.5 \times 10^3$ CFU mL <sup>-1</sup> Visual $2.39 \times 10^2$ CFU mL <sup>-1</sup> FL	Specific, Optimization of QD amount and MNP, Silica layer have NH <sub>2</sub> functional group and separated QDs from MNPs and decrease the	Milk	[365]

AuNPs	AuNPs <sub>c</sub> decorated with Raman label, AuNPs <sub>c</sub> 21.56± 4.52, for Antibody interaction pH 9 is optimum	A <i>E. coli ATCC 35218</i> b	LFIA/ SERS/ CL	$10^3\text{-}10^8$ CFU mL <sup>-1</sup> $10^2\text{-}10^6$ CFU mL <sup>-1</sup> SERS	78CO 45 SERS CFU mL <sup>-1</sup>	10 mm Pore size had a slightly higher intensity. DTNB better than Glutathione as Raman label	Urine Blood	[366]
AuNPs	AuNPs 20nm	A <i>E. coli O157:H7</i> , b <i>S. Typhimurium</i> , <i>S. aureus</i> , <i>Bacillus cereus</i>	LFIA	$10^1\text{-}10^6$ CFU mL <sup>-1</sup>	$10^5\text{-}10^6, 10^5$ CFU mL <sup>-1</sup> <i>E.coliO157:H7</i> , <i>S.Typhimurium</i> , <i>S. aureus</i> , <i>Bacillus cereus</i> CFU mL <sup>-1</sup>	Rotary type device $1.87 \times 10^4$ CFU of <i>E. coli O157: H7</i> $1.47 \times 10^4$ of <i>S. Typhimurium</i> , 1 CFU g <sup>-1</sup> <i>E. Coli O157: H7</i> and <i>S. Typhimurium</i> after 6 h preconcentration	Lettuce	[367]
UCP(NaYF4:Yb <sup>3+</sup> ),Er <sup>3+</sup>	Stable optical property	A <i>Francisella tularensis</i> b	LFIA	$10^4\text{-}10^8$ CFU mL <sup>-1</sup>	$10^3$ CFU mL <sup>-1</sup>	Selective, pH 2-13, high ion strengths , high viscosities, high concentrations of bio macromolecules showed little influence	Seven powders Eight viscera	[368]
AuNPs	AuNPs <sub>c</sub>	A <i>Salmonella serogroups</i> b	LFIA/ CO	-	-	Arrangement of five <i>Salmonella</i> serogroups. Three test line	Feces Anal swab	[369]
Hierarchical flowerlike AuNPs	20 nm AuNPs as seeds & polyelectrolyte PDDA as polymer capping agent to synthesize hierarchical flowerlike Au particles, AuNPs 20,40 nm	A <i>E. coli O157: H7</i> b	LFIA/ CO	$10^3\text{-}10^7$ CFU mL <sup>-1</sup>	$10^3$ CFU mL <sup>-1</sup>	Comparing 5 different nanoparticles, diverse shaped AuNPs showed better sensitivity 2 order of magnitude better than spherical AuNPs 20 nm	-	[370]

AuNPs	AuNPs <sub>C</sub> 18 nm	A <i>E. coli</i> O157: H7 b	pLFIA	100-600 CFU mL <sup>-1</sup>	100 CFU mL <sup>-1</sup>	Colorimetric signal improved by utilizing liposome encapsulating polyethyleneimine to activate the aggregation of gold nanoparticles 1000 fold better sensitivity, 600 CFU mL <sup>-1</sup> <i>E. coli</i> O157: H7 in cranberry juice	Cranberry juice	[179]
QD	-	A <i>E. coli</i> p	FLFA	-	15-150 Viable cell	Specific, Test sequence	7 Big aptamer	- [371]
AuNPs Magnetic bead	AuNPs 15 nm citrate	A <i>E. coli</i> O157: H7 p	SDA/L FA	10-10 <sup>6</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>		Milk powder, Water, Apple juice	[186]
AuNPs		A <i>Vibrio parahaemolyticus</i> p	SDA/L FA	10-10 <sup>7</sup> CFU mL <sup>-1</sup>	10 CFU mL <sup>-1</sup>	Selective, 200 CFU mL <sup>-1</sup> in Oyster	Oyster	[372]
Au <sup>MBA</sup> @Ag core-shell nanoparticles	Au <sup>MBA</sup> @Ag core the diameter of 23.58 nm shell thickness 8.48 adding MBA solution to the AuNPs solution followed by ascorbic acid reduction of silver nitrate	DNA probe <i>L. monocytogenes &amp; S. enterica</i>	RPA/L FA/SE RS	1.9 × 10 <sup>1</sup> -1.9 × 10 <sup>6</sup> 2.7 × 10 <sup>1</sup> -2.7 × 10 <sup>6</sup> CFU mL <sup>-1</sup> <i>L. monocytogenes &amp; S. enterica</i>	19 & 27 for CFU mL <sup>-1</sup> <i>L. monocytogenes &amp; S. enterica</i>	Specific	Milk, Chicken breast, Beef	[188]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 30 – 40 nm	DNA probe <i>HIV-1 viruses</i>	SERS/ LFA	10 <sup>-8</sup> -10 <sup>-4</sup> gL <sup>-1</sup>	8 × 10 <sup>9</sup> mL <sup>-1</sup> , Naked eye 8 × 10 <sup>8</sup> gL <sup>-1</sup>	Specific	-	[373]
Mesoporous silica nano-particles	Silica nanoparticle prepared with sol-gel method hexagonal pore structure 2.4 nm mesopores	A <i>Listeria monocytogenes</i> p	LFA/ Ap gated releas e	3 × 10 <sup>1</sup> -3 × 10 <sup>6</sup> CFU mL <sup>-1</sup>	53 cells mL <sup>-1</sup>	Selective	Minced Chicken meat	[187]
AuNPs <sub>C</sub>	AuNPs <sub>C</sub> 20 nm, Polystyrene beads -Au- PEI particles	A <i>E. coli</i> O157: H7 p	LFA/ digital camera	10-10 <sup>3</sup> CFU mL <sup>-1</sup>	25 CFU mL <sup>-1</sup> , 253 CFU mL <sup>-1</sup> Naked eye, 233 CFU mL <sup>-1</sup> Ground beef	Specific, Novel Carboxyl functionalized DNA aptamers to effectively graft the aptamer on the label particles surface, Inkjet printer fabrication for patterning of aptameric ink	Ground beef	[374]

nanoparticle in the two test lines containing specific capturing probes while the SERS signal of test lines was measured for quantitative detection of bacteria. SERS signal was shown to be linear with target concentration in the  $1.9 \times 10^1$ – $1.9 \times 10^6$  and  $2.7 \times 10^1$ – $2.7 \times 10^6$  interval for *L. monocytogenes* and *S. enterica* respectively [188]. More examples of lateral flow based biosensor are available in Table 6.

#### 4. Conclusion and future prospective

In this review, various nanomaterials and their application in different detection systems for infectious agents covered. Nanomaterials as magnetic nanoparticles alone and in combination with Ag or Au coatings, bimetallic nanoparticles, peroxidase-like nanomaterials, and newly designed nanomaterials have been described and their widespread application in electrochemical and optical biosensing discussed. Applications as separation materials, materials for improving conductivity, and elements for biomolecules grafting have been described. Besides, their application in colorimetric, fluorometric, and SERS detection methods was also reported. Finally, we demonstrated the advantage of the usage of these materials to enhance the sensitivity of the lateral flow strip test. Although some authors illustrated comparative data to highlight the benefits of the usage of nanomaterials, there is still the need for more investigation to understand the real effect and advantages of exploiting nanomaterials in various sensing applications. In the future, we expect novel functional biomimetic and bimetallic nanomaterial or bionanocomposite with fascinating features will be introduced. Moreover, innovative designs strategies with the help of biochemical pathways to shorten the analysis time, to obtain the results of the molecular techniques such as PCR and LAMP and to produce efficient and rapid POC devices to be used in the field will report. Nanoenzyme material may replace natural recognition receptors to produce more durable biosensors. One of the biggest challenge for the nanomaterials are their work in various real-samples matrices such as blood, serum, urine or stool in biomedical or various food matrices or for example sewage water in case of environmental samples. Therefore we expect a lot of attention will be directed towards the development of more efficient nanomaterials characteristics with minimum non-specific

binding and low false positive signals. We also expect much attention will be paid in the future towards the development of multifunctional nanomaterials where it integrates for example specific analyte fishing or preconcentration tool as well as sensing platform. Despite the numerous proposed biosensors, there is still a crucial need for developing new biosensors for on-site monitoring of infectious diseases to prevent epidemic and economic loss.

#### Declaration of competing interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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