

GRAPHENE-BASED DIELECTROPHORESIS SENSOR AND METHOD

CROSS-REFERENCE TO RELATED APPLICATIONS

This application claims priority to U.S. Application Serial No. 62/680,777, filed
5 on June 5, 2018. The disclosure of the prior application is considered part of the
disclosure of this application, and is incorporated in its entirety into this application.

BACKGROUND

According to the World Health Organization, 550 million people become ill from
foodborne enteric diseases, and 230,000 people die from them each year. Infections from
10 *Salmonella* spp. and *Escherichia coli* (*E. coli*) rank among the most common of these
enteric diseases. *Salmonella* spp. is a bacterium found in animals that, if ingested, can
cause serious health problems. Such problems are particularly serious in people with high
risk for complications such as young children, older adults and people with weakened
immune systems. Similarly, *E. coli* is a coliform bacterium that can cause sickness or
15 even death in humans, depending on the strain of the bacteria and the susceptibility of the
person. In Minnesota and across the country, *Salmonella* spp. and *E. coli* outbreaks
continue to be an ongoing threat to public health.

Commonly used test methods for pathogen bacterial DNA involve culture and/or
polymerase chain reaction (PCR) analysis, which is a laboratory technique used to
20 amplify a target DNA molecule using a series of repeated amplification sequences. PCR
has excellent sensitivity, but requires dozens of cycles to determine the presence or
absence of a particular species, is both time-consuming and costly, and often requires an
initial enrichment culture step. Increasingly rapid and sensitive methods to detect
pathogenic organisms in food products or clinical samples are being developed. In
25 particular, methods to detect *Salmonella* spp. rapidly have been pursued because of the
importance of this bacteria as a foodborne pathogen, recent advances have further
improved the speed and sensitivity of detection using other methods, but these techniques
still fall short of the speed and sensitivity required for hand-held systems.

A particularly promising method of DNA detection of hand-held systems involves the use of graphene-based sensors, and numerous reports in the literature have shown that graphene can provide an excellent alternative method to selectively sense DNA. Graphene is a two-dimensional sheet of sp^2 -bonded carbon, which can be realized in monolayer form either through mechanical exfoliation, or through growth on copper using chemical vapor deposition (CVD).

Due to its monolayer nature, graphene can provide excellent sensitivity to adsorbed biomolecules, and can also be functionalized to provide selectively to chemically-active targets.

Conventional graphene sensors utilize simple diffusion processes to attract particles to the surface. This can be slow. On the other hand, dielectrophoresis (DEP) can be used to attract particles to a region of high electric field gradient, but does not provide a convenient means of selective sensing. In this invention, a sensor is described that solves the problem of how to achieve selectivity within a DEP-based sensors by the use of graphene which is functionalized with a probe material on the surface. It also provides a way to separate the DEP-attraction bias voltages, which are relatively large, and the sensing bias voltages, which are often smaller.

SUMMARY

In a first aspect, the present disclosure provides a dielectrophoretic (DEP) sensor, comprising: a graphene electrode adjacent a channel for confining a target particle in a liquid; a surface probe attached to a surface of the graphene electrode, the surface probe having a selective reaction with the target particle; and a voltage source electrically connected to the graphene electrode and configured to (i) apply a voltage to the graphene electrode to cause DEP trapping of the target particle at the graphene electrode, and (ii) apply a measurement signal to the graphene electrode, the measurement signal being sufficient to readout a response from the DEP sensor indicative of a presence of the target particle.

In a second aspect, the present disclosure provides a readout method for a dielectrophoretic (DEP) sensor comprising a graphene electrode with a surface probe attached to a surface of the graphene electrode, the surface probe having a selective

reaction with a target particle, the readout method comprising: applying an AC excitation to the graphene electrode sufficient to dielectrophoretically attract the target particle toward the graphene electrode; and applying a measurement signal to the graphene electrode, the measurement signal being sufficient to readout a response from the DEP sensor indicative of a presence of the target particle.

In certain embodiments, the disclosure features a graphene-edge dielectrophoretic (DEP) sensor that utilizes surface functionalization to achieve specificity and selective sensing. In some embodiments, the disclosure features a methodology by which to detect the selective response of the sensor.

In some embodiments, the graphene-based DEP sensor utilizes surface functionalization on the graphene or adjacent surfaces (or substrates) to achieve selectivity. The graphene-based DEP sensor can use a time-varying excitation for DEP attraction that enables the sensor response to be read out independent of the DEP AC excitation.

Among other advantages, embodiments can achieve selectivity to a range of target molecules or biological particles using a DEP-based graphene sensor. Embodiments can

also enable reading out the sensor response in the presence of a relatively large AC excitation meant to attract particles to the graphene edge.

The details of one or more embodiments of the disclosure are set forth in the accompanying drawings and the description below. Other features, objects, and advantages of the disclosure will be apparent from the description and drawings, and from the claims.

DESCRIPTION OF DRAWINGS

FIGS. 1a – 1c are schematic diagrams showing a top view of a resistance-based DEP sensor. Fig. 1a shows the sensor with the attached probe on the graphene surface, in a solution containing the target. Fig. 1b shows the sensor with the targets attracted to the graphene edges using DEP via an applied AC bias. Fig. 1c shows the targets bonding to nearby probes after the DEP excitation is removed, enabling sensor readout.

FIGS. 2a – 2c are schematic diagrams showing a top view of a capacitance-based DEP sensor. Fig. 2a shows the sensor with the attached probe on the graphene surface, in a solution containing the target. Fig. 2b shows the sensor with the targets attracted to the graphene edges using DEP via an applied AC bias. Fig. 2c shows the targets bonding to nearby probes after the DEP excitation is removed, enabling sensor readout.

FIG. 3 shows plots of the applied voltage signal and anticipated response for a resistance-based DEP sensor.

FIG. 4 shows plots of the applied voltage signal and anticipated response for a capacitance-based DEP sensor.

FIGS. 5a and 5b show plots of an anticipated response of a capacitance-based DEP sensor with varied AC bias.

FIGS. 6a – 6e illustrate a graphene-edge DEP tweezer fabrication process. FIG. 6a shows gate oxide recess and gate metal patterning. FIG. 6b shows graphene transfer. FIG. 6c shows graphene patterning into trapping segments. FIG. 6d shows contact metallization and lift off. FIG. 6e shows the surface functionalization with a probe object.

Like reference symbols in the various drawings indicate like elements.

DETAILED DESCRIPTION

FIG. 1a shows an embodiment of a graphene-edge dielectrophoretic sensor 105. The particular sensor operates in the resistive sensing mode as we have described previously in Provisional Application No. 62/521,096, entitled “Electrodes formed from 2D materials for dielectrophoresis and systems and methods for utilizing the same,” filed on June 16, 2017, and in U.S. 2018/0361400, filed Jun. 18, 2018, the entire contents both of which are hereby incorporated by reference.

DEP sensor 105 includes the first fingered electrode 113, a graphene electrode 182, and contacts 184. The contacts 184 are electrically coupled to the graphene electrode 182. The DEP sensor 105 is a three-terminal device, in which the first fingered electrode 113 serves as a gate electrode, and the two contacts 184 serves as the source and drain terminals. The graphene electrode 182 is arranged in a serpentine shape to increase crossing sites with the first fingered electrode 113. Increasing the crossing sites leads to increased number of edges where particles can be trapped. For example, widths of the serpentine strips of the graphene electrode 180 can range from 100 nm to 5 μm , and widths of the fingers of the first fingered electrode 113 can range from 0.1 μm to 5 μm .

The graphene electrode 182 is electrically isolated from the first fingered electrode 113. For example, an insulating layer similar to the insulating layer 130 may be arranged between the first fingered electrode 113 and the graphene electrode 182. As such, an example stack-up of the DEP sensor 105 may include, in the order as listed, the first fingered electrode 113, the insulating layer, and the graphene electrode 182. The insulating layer may act as the “gate oxide” of the 3 terminal FET device formed by the graphene electrode 182 and the first fingered electrode 113.

The contacts 184 may be formed from various conductive materials. Examples of conductive materials include metals, silicides, or alloys, such as Gold, Palladium, Platinum, Tungsten, Chromium, Titanium, Iridium, Molybdenum, Aluminum, or Copper. In some implementations, different layers of conductive materials can be stacked to form the contacts 184.

The DEP sensor 105 can be electrically measured by applying a DC voltage, an AC voltage, or both (collectively referred to as “measurement voltage”) between the two ends of the graphene electrode 182 through the contacts 184, and measuring the resulting

drain current (ID) while applying an AC voltage ("trapping voltage") between the first fingered electrode 113 ("gate electrode") and the graphene electrode 182 to induce DEP trapping. The trapping voltage can be applied through a voltage source 186 and the measurement voltage can be applied through a source meter 188 configured to apply a voltage and measure corresponding current. Initially, a trapping voltage is applied between the gate electrode 113 and the graphene electrode 182 at a frequency that does not correspond to DEP trapping (e.g., 1 MHz) for a particular type of particles, and no particles are trapped.

The measurement voltage is typically smaller than the trapping voltage to mitigate a voltage drop across the two terminals of the graphene electrode 184 caused by the flow of the drain current ID. For example, the measurement voltage can be 10 to 100 times smaller than the amplitude of the trapping voltage.

In some implementations, the trapping voltage can have a DC bias in addition to the AC voltage. The DC bias can be used, for example, to bias the graphene electrode 182 to control its conductivity.

As shown in FIG. 1a, a surface probe 101 is attached to the graphene electrode 182, where the probe is designed to have a selective reaction with a target 102. The probe and target can feature of any complementary pair of molecules, proteins, aptamers, antibodies, lipids, vesicles, cell-derived particles, functionalized nanoparticles or whole cells that have the property that the probe can be attached to the surface of the graphene, and that the target is a polarizable particle such that it can be attracted to the graphene surface using DEP and can selectively attach to the surface probe on the graphene.

An example of such a probe/target pair is single-stranded DNA molecules with complementary matching. Another example is functionalization of graphene using biotin or biotin-labeled proteins for the detection of streptavidin targets.

Selective attachment of the target to the probe can be detected using the sensor. This can be achieved using either the capacitive or resistive sensing means described previously in Provisional Application No. 62/521,096 and U.S. 2018/0361400 using a graphene-edge DEP geometry. The sensing can be achieved using an "attract-and-listen" mode, where the target particles are first attracted to the graphene surface using a DEP

excitation, and then the excitation is turned off and the graphene sensor response is evaluated using a smaller signal.

A diagram of this mode is illustrated in FIGS. 1a-1c which show DNA detection, where prior to the DEP attraction pulse, the target DNA is attracted to the regions of the graphene where the edges cross a gate edge. After release of the DEP pulse, as shown in FIG. 1c, the target DNA can attach to the probe DNA on the graphene surface nearby thus creating an electrical response. This response can be achieved using the resistive configuration shown in FIGS. 1a – 1c or in a capacitive configuration as illustrated in FIGS. 2a – 2c.

FIG. 2F shows a top view of a graphene varactor-integrated DEP device 204. The DEP device 204 includes a first fingered electrode 213, a second fingered electrode 233, and a graphene electrode 280. The graphene electrode 280 is electrically coupled to the second fingered electrode 233. The graphene electrode 280 has been subdivided into several smaller fingers to increase the number of edges and crossings of the graphene electrode 280 over the first finger electrode 213, where particles can be trapped. For example, the width of the subdivided fingers can range from 100 nm to 5 μ m.

The graphene electrode 280 is electrically isolated from the first fingered electrode 213. For example, an insulating layer similar to the insulating layer 130 may be arranged between the first fingered electrode 213 and the graphene electrode 280. As such, an example stack-up of the DEP device 204 may include, in the order as listed, the first fingered electrode 213, the insulating layer, the graphene electrode 280, and the second fingered electrode 233. Further, other layers of materials may be present between the first fingered electrode 213, the insulating layer, and the graphene electrode 280.

The first and second fingered electrodes 213 and 233 may be formed from various conductive materials. Examples of conductive materials include metals, silicides, or alloys, such as Gold, Palladium, Platinum, Tungsten, Chromium, Titanium, Iridium, Molybdenum, Aluminum, or Copper. In some implementations, different layers of conductive materials can be stacked to form the first and second electrodes 213 and 233. Such stacking may be beneficial for improving adhesion between the metal and the insulating layer or the graphene, and for improving the quality of electrical contact to the graphene electrode 280.

Initially, an AC bias is applied between a the first fingered electrode 213 (“gate electrode”) and the second fingered electrode 233 coupled to the graphene electrode 280 at a frequency that does not correspond to DEP trapping (e.g., 1 MHz) for a particular type of particles, and no particles are trapped as a result. The AC bias can be applied
5 through a voltage source 216.

The voltages applied (lower plot) and the anticipated response (upper plot) are shown in FIG. 3 for the resistive sensing mode. Here, the current can be readout in this device by applying a DC drain-to-source voltage, V_{DS} , and a constant gate voltage V_{GS} . When the DNA is applied to the sample, the response is typically slow. However, if a
10 larger amplitude AC V_{GS} signal is applied, DEP allows attraction of the DNA to the sample edges, accelerating the device response. Once the AC signal is removed the sensor reads a change in current, associated with trapping the target molecules to the surface. This readout could also be achieved by sweeping the gate voltage and monitoring the current to determine the gate voltage, the Dirac voltage, at which a
15 minimum current occurs.

In FIG. 4, a sensing scenario for a capacitive-based sensor is shown. Here, the sensor is read out in a simple two-terminal configuration using a small AC gate bias. Once again, to accelerate the device response, a larger AC bias can be applied and then reduced again to read out the sensor capacitance response. Similar to the resistive sensor,
20 the readout could also be achieved by sweeping the gate voltage and monitoring the Dirac voltage, which is the gate voltage at which the capacitance is minimum.

As shown in FIG. 5, in some embodiments, this scenario is repeated and utilized for different concentrations of the target analyte, with the benefit of speeding the response compared to allowing the analytes to move to the surface through simple
25 diffusive processes. This also differs from an alternative mode of sensing, where the DEP signal is applied continuously and the sensor response is read out concurrently with the DEP signal. A similar pulse sequence could be utilized to perform negative DEP, where unwanted particles are repelled from the surface after the sensor is exposed to the target-containing fluid. The sensing modality would be similar, except the frequency of the
30 excitation would be such that certain particles in solution would be repelled.

In general, the sensors and techniques disclosed above can be implemented in a variety of applications. For example, they can be deployed in point of care sensors. In some embodiments, the embodiments are deployed in DNA-based bar-code sensors.

5 Prophetic Example 1: fabrication and trapping verification of sensors.

DEP-based sensors are fabricated and their trapping capability validated as follows. The existing graphene variable capacitor (or “varactor”) fabrication process (*see, e.g.,* M. A. Ebrish, H. Shao, and S. J. Koester, “Operation of multi-finger graphene quantum capacitance varactors using planarized local bottom gate electrodes,” Appl. Phys. Lett. 100, 143102 (2012).) is modified to make it suitable for trapping large densities of particles near the graphene surface.

An example fabrication process for the graphene-edge DEP sensors is shown in FIGS. 6a – 6e. The device fabrication starts with a silicon wafer upon which thick SiO₂ is grown by thermal oxidation. Next, a gate contact electrode is patterned using optical lithography and the oxide recessed to a depth of ~50 nm and then gate metal (usually Ti/Pd) is deposited and lifted off to form a quasi-planar electrode (FIG. 6a). After the gate metallization, thin HfO₂ is deposited (usually ~ 7-10 nm) on the entire sample using atomic-layer deposition (ALD). Next, chemical vapor deposited (CVD) graphene is transferred onto the sample using a PMMA handle layer, where the PMMA is subsequently removed using a solvent clean and vacuum annealing (FIG. 6b).

Next, the graphene is patterned into a series of narrow strips, creating a greater number of edges where molecules can be trapped. Using optical lithography, the strips are as small as 2 μm wide with 4 μm pitch. Using electron-beam lithography, the strips are as narrow as 100 nm with 200 nm pitch. A schematic of the patterned graphene is shown in FIG. 6c. Next, Cr/Au (10/100 nm) metallization is patterned to form Ohmic contacts to the graphene as shown in FIG. 6d. The device fabrication can also include a thick (several micron) isolation layer such as SU-8 which is patterned and opened only above the active device area to form a micro-volume for fluid confinement during testing. Finally, functionalization is applied to attach probes to the graphene surface, as shown in FIG. 6e.

Different device geometries can be explored to determine an optimal structure to realize simultaneous trapping and sensing. There are several factors in this design optimization. These can include: (1) the number of edges necessary in order to achieve sufficiently effective trapping; (2) capacitance or resistive sensing; and (3) the optimal sequence of excitations to attract and sense the DNA fragments.

The sensor can be tested using filtered DNA samples in 1 mM KCl buffer (*see, e.g.,* A. Barik, Y. Zhang, R. Grassi, B. P. Nadappuram, J. B. Edel, T. Low, S. J. Koester, and S.-H. Oh, “Graphene-edge dielectrophoretic tweezers for trapping of biomolecules,” *Nat. Commun.* 8, 1867 (2017).), however more complex media with additional particles such as cell membranes and protein residue can be investigated. For food-borne pathogen detection, the DNA samples could include fragments of *Salmonella typhimurium*. The fragments vary in length from 1 kb to 100 bp based upon the type of PCR method (gel-based, cybergreen, or real-time) used to obtain the samples.

Prophetic Example 2: DNA probe attachment to graphene and selective DNA sensing for *Salmonella typhimurium* and *E. coli* using graphene-edge DEP tweezers.

This example demonstrates two main elements: (1) surface functionalization of probe DNA on graphene and (2) sensing verification using the device structure shown in FIG. 6(d).

1-pyrenebutyric acid N-hydroxysuccinimide ester (PBASE) is attached to the graphene surface. The pyrene end of the PBASE attaches to the graphene through π - π interactions, while the succinimide portion extends outward from the graphene enabling bonding to the probe DNA. In order to attach the probe, the 5' end of the DNA is modified to include an amine group. This modified DNA is exposed to the PBASE-covered graphene in solution, and allowed to crosslink with the succinimide to form a stable functionalization layer. In general, the functionalization procedure is achieved by soaking the entire substrate in a probe-DNA-containing solution such as acetonitrile, though this process can readily be modified to achieve locally functionalized devices on the same chip using nozzle-based or 3D printing, thus enabling multiplexed sensing capability.

The attachment of the probe DNA to the graphene surface can first be verified on blanket graphene layers using atomic force microscopy (AFM) and Raman spectroscopy. Functionalization molecules on graphene can be verified using Raman spectroscopy, both by identifying peaks specific to the functionalization species as well as analyzing shifts in the G and 2D peaks that demonstrate systematic changes in the electron or hole concentration in the graphene. After functionalization has been verified using these techniques, the DNA probes are attached to graphene on a sensor device structure. In these structures, the probe DNA attachment is further verified by analyzing the shift in the Dirac voltage. It is believed that Dirac point shifts are a reliable indicator of surface functionalization attachment as has been demonstrated using pyrene boronic acid functionalization (*see, e.g.*, Reference Y. Zhang, R. Ma, X. Zhen, Y. C. Kudva, P. Bühlmann, and S. J. Koester, "Capacitive sensing of glucose in electrolytes using graphene quantum capacitance varactors," ACS Appl. Mater. Interfaces 9, 38863–38869 (2017)). Here, the graphene functionalization can be verified using increasing concentrations of the probe-DNA containing solutions and the Dirac point shift monitored as a function of concentration. This verification is important, since residues from device processing could be present and comparison of the equivalent doping results from the bare graphene via Raman and the device graphene from electrical tests provide an important validation for subsequent sensing.

DEP-enhanced sensing of the graphene devices is verified as follows. Medium-to-high conductivity KCl solution are utilized with concentration roughly $0.1\times$ of standard phosphate buffered saline (PBS). In the initial tests, 100-1000 bp DNA fragments extracted from *E. coli* and *Salmonella typhimurium* PCR are evaluated at concentrations ranging from 1 nM down to 1 fM. The probe DNA consists of single-stranded DNA and the target DNA consists of single-stranded DNA with complementary base pair matching to the probe.

The samples are tested using a flow cell geometry. In the testing procedure, a solution containing the target DNA is cycled over the exposed region of the prefunctionalized sensor. The flow-cell geometry allows a single device to be tested at varying DNA concentrations.

Parasitic effects caused by the conductivity of the buffer solution are minimized due to the SU-8 layer that isolates the extrinsic region of the sensor and only allows the active sensor region to interact with the test solution. However, if such issues persist, additional dilution of the sensing solution should be possible. A test chip that allows
5 testing of the devices using a standard probe station is used, where the probes are located far from the active device region in the test liquid. In addition, all of the sensing experiments utilize differential sensing, where a “blank” sensor (functionalized with pyrene only) is utilized as the control device. By subtracting signals from the electrode without receptors, background signals, drift issues, and temperature variations can be
10 rejected. Such differential sensing may be important in DEP where heating effects are possible due to the fluid motion induced by the AC excitation.

Two main sensing protocols are described, based upon the two types of sensors geometries shown in FIGS. 1a-1c and 2a-2c. The first is the resistance-based geometry shown in FIGS. 1a-1c. In this geometry, the graphene is patterned in a shape where the
15 graphene crosses a multi-finger buried electrode several times. The graphene can be arranged in a serpentine geometry to increase the number of gate edge crossings within a specific area. In this geometry, a small DC or AC bias (on the order of 100 mV) is applied between the ends of the graphene serpentine. The sensor response is determined by monitoring the relative drain current change ($\Delta I_D/I_D$) at fixed values of V_{DS} vs. time for
20 various DNA concentrations. The sensing can be monitored without DEP, by simply setting the gate voltage, V_{GS} , to a fixed value and monitoring $\Delta I_D/I_D$ vs. time. This response time is set the baseline with which to compare the DEP-enhanced sensing and is limited by the diffusion time and subsequent hybridization of the target DNA at the sensor surface.

25 After this baseline is established, the graphene-edge DEP sensing is evaluated. For instance, at each concentration, the DC response is monitored for 5 minutes to establish a baseline, and then an AC excitation is added to the gate electrode for a prescribed time (e.g., 5 minutes), and removed and the DC response monitored again. The AC voltage is centered about zero (no DC component) and the peak-to-peak value
30 and frequency is determined based upon trapping dependent studies. In this geometry, it may be important to keep V_{DS} much smaller than the AC peak-to-peak signal during

trapping, to ensure the trapping efficiency does not change along the length of the sensors.

The second device geometry that is evaluated is the varactor geometry shown in FIGS. 2a-2c. The geometry operates in a parallel configuration and so small breaks in the graphene at the gate edges should not affect performance. The device also operates at $V_{DS} = 0$, and thus does not have the problem of sensing variations at different trapping sites. The varactor configuration has the advantage that capacitance measurements are already performed at high frequencies, and thus only the amplitude of the excitation needs to be changed in order to trap DNA.

Finally, the selectivity of the sensors against interfering particles is evaluated in order to mimic the effect of sensing in realistic biological cultures. Since DNA molecules have higher electrical polarizability than proteins and lipids, it is expected that an optimal DEP frequency range and voltage can be found to concentrate DNA molecules selectively while rejecting unwanted interferants and debris.

A number of embodiments have been described. Nevertheless, it will be understood that various modifications may be made without departing from the spirit and scope of the invention. Accordingly, other embodiments are within the scope of the following claims.

WHAT IS CLAIMED IS:

1. A dielectrophoretic (DEP) sensor, comprising:
a graphene electrode adjacent a channel for confining a target particle in a liquid;
a surface probe attached to a surface of the graphene electrode, the surface probe having a selective reaction with the target particle; and
a voltage source electrically connected to the graphene electrode and configured to (i) apply a voltage to the graphene electrode to cause DEP trapping of the target particle at the graphene electrode, and (ii) apply a measurement signal to the graphene electrode, the measurement signal being sufficient to readout a response from the DEP sensor indicative of a presence of the target particle.
2. The DEP sensor of claim 1, wherein the surface probe and target particle are a complementary pair selected from the group consisting of molecules, proteins, aptamers, antibodies, lipids, vesicles, cell-derived particles, functionalized nanoparticles and whole cells.
3. The DEP sensor of claim 1 or 2, wherein the graphene electrode comprises a plurality of strips.
4. The DEP sensor of claim 3, wherein the strips are parallel to each other.
5. The DEP sensor of any one of the preceding claims, further comprising a second electrode overlapping with the graphene electrode and electrically connected with the voltage source.
6. The DEP sensor of claim 5, further comprising a source meter in electrical communication with the graphene electrode and the second electrode.
7. The DEP sensor of any one of the preceding claims, wherein the DEP sensor is a resistance-based sensor.

8. The DEP sensor of any one of the preceding claims, wherein the DEP sensor is a capacitance based sensor.

9. A readout method for a dielectrophoretic (DEP) sensor comprising a graphene electrode with a surface probe attached to a surface of the graphene electrode, the surface probe having a selective reaction with a target particle, the readout method comprising:

applying an AC excitation to the graphene electrode sufficient to dielectrophoretically attract the target particle toward the graphene electrode; and

applying a measurement signal to the graphene electrode, the measurement signal being sufficient to readout a response from the DEP sensor indicative of a presence of the target particle.

10. The readout method of claim 9, wherein the measurement signal is a DC signal.

11. The readout method of claim 9, wherein the measurement signal is an AC signal.

12. The readout method of any one of claims 9-11, wherein an amplitude of the AC excitation is larger than an amplitude of the measurement signal.

13. The readout method of any one of claims 9-12, wherein the readout is based on a change of capacitance measured by the sensor.

14. The readout method of any one of claims 9-12, wherein the readout is based on a change of current measured by the sensor.

15. The readout method of any one of claims 9-14, wherein the surface probe and target particle are a complementary pair selected from the group consisting of molecules, proteins, aptamers, antibodies, lipids, vesicles, cell-derived particles, functionalized nanoparticles and whole cells.