



Design and analysis of a multi-modal refractive index plasmonic biosensor based on split ring resonator for detection of the various cancer cells

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Received: 26 July 2024 / Accepted: 16 August 2024

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Abstract

This paper introduces an innovative optical biosensor designed for multimode refractive index (MMRI) detection, employing split ring resonator (SRR) technology for enhanced sensitivity in cancer cell analysis. The proposed biosensor comprises two concentric SRRs, one resembling a cube and the other a cylinder, each incorporating an air gap. Rigorous analysis and optimization of dimensional parameters were conducted using the finite difference time domain (FDTD) numerical solution method. The simulation results demonstrated that the biosensor achieves exceptional sensitivity and an optimal figure of merit (FOM). Specifically, the first mode of the biosensor exhibited a sensitivity exceeding 80 nm/RIU and an FOM surpassing 20 RIU⁻¹, while the second mode demonstrated a sensitivity of 570 nm/RIU and an FOM exceeding 4 RIU⁻¹. These results indicate that the first mode benefits from a higher FOM, enhancing precision and accuracy, while the second mode benefits from high sensitivity, enabling the detection of minute refractive index changes. With a primary focus on early cancer cell detection and timely treatment initiation to address the global cancer mortality burden, the biosensor's high-quality factor (Q-factor) and FOM promise significant advancements in cancer diagnostics and therapeutic interventions.

Keywords Optical biosensor · Plasmonic · Split ring resonator · Cancer cells · Multimode refractive index · Sensitivity

1 Introduction

Cancer is one of the leading causes of death globally, according to scientific studies, and researchers have focused their attention on its early detection because it is crucial for initiating an effective treatment regimen (Bray et al. 2018; Yang et al. 2019). There are numerous methods available for the diagnosis of cancer, with tissue biopsy being the prevailing technique utilized globally. In recent years many advances in liquid

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biopsy technologies have been investigated to detect cancer biomarker (Soda et al. 2022). Nevertheless, in numerous instances, patients who undergo a tissue biopsy may encounter potential risks, including the sampling of tumors situated near vital organs or lesions located in challenging regions of the brain (Robinson et al. 2024). As a result, the exploration for minimally invasive biomarkers plays a critical role in enabling early detection technology with consistent monitoring in tandem with cancer therapy. In recent times, a method called liquid biopsy has been introduced in the medical field to unveil the molecular composition of solid tumors using blood samples. Liquid biopsy has become a widely used medical technique for the identification and surveillance of cancer tumors by detecting various biomarkers such as circulating tumor cells (CTCs), circulating tumor nucleic acids (ctDNA, ctNAs, microRNA), and extracellular vesicles (EVs). Given that these biomarkers can be found in bodily fluids like blood, oral saliva, and urine, liquid biopsy offers a less invasive alternative to tissue biopsy, resulting in reduced pain and lower risk of infection (Wan et al. 2017; Ignatiadis et al. 2021). In recent years, sensor-integrated microfluidic approaches have shown significant promise for liquid biopsy applications in the early detection of cancer, offering enhanced sensitivity and specificity in detecting cancer biomarkers (Sierra et al. 2020). Dielectric grating with a box-like resonance shape as a novel ultra-compact photonic biosensor with a tailored resonance shape has presented in Toma et al. (2024).

The refractive index, electrical permeability, magnetic permeability, and various optical characteristics inherent to biological materials offer valuable insights into their elemental and chemical composition. Consequently, the manipulation of these optical properties in biological samples, coupled with the utilization of optical biosensors, constitutes a critical aspect of optical detection methodologies for analyzing such samples, including cancer cells (Mårtensson Jönsson 2015; Liu et al. 2016). The optical biosensors that have been suggested demonstrate that the identification of extracellular vesicles (EVs) relies on appropriate bio detection components and refractive index (RI) sensitivity. This is due to the fact that the presence of EVs induces a modification in the RI of the specimen. Consequently, it is crucial to establish a photonic sensor architecture that possesses a heightened RI sensitivity in order to achieve real-time, label-free, and effective cancer detection through EV biomarkers (Saha et al. 2022). The design and construction of sensor devices necessitate research and study across diverse disciplines, encompassing engineering sciences, physics, biology, and chemistry. This multidisciplinary approach is essential for the development of sensor technologies that can effectively address complex challenges and meet the demands of diverse applications (Mehrotra 2016; Coulet and Blum 2019). Recent research discussed in Mondal et al. (2024) underscores the significant role of plasmonic biosensors in digital healthcare applications. These biosensors are instrumental in facilitating early disease diagnosis, biomarker monitoring, pathogen detection, and drug screening. By integrating these sensors with digital platforms and smartphone interfaces, real-time data collection, analysis, and transmission are made possible, thereby promoting the transition towards interconnected and readily available healthcare solutions. Jafrasteh et al. introduced a plasmonic sensor designed for the detection of cancer and undertook comprehensive research to assess food contaminants utilizing machine learning and artificial intelligence. Furthermore, they demonstrated that the adverse impacts of environmental pollution on human health can be mitigated through the application of artificial intelligence and machine learning techniques (Jafrasteh et al. 2023). Salehnezhad et al. developed and simulated a highly sensitive plasmonic nanosensor that incorporates monolayer MoS₂ and graphene, achieving a sensitivity of 192°/RIU. This innovative sensor is adept at detecting antibodies with refractive indices that closely resemble those of blood groups (Salehnezhad

et al. 2023). Furthermore, based on the findings of Farmani et al., a label-free nano sensor utilizing graphene and structured surface plasmon resonance was developed, exhibiting a sensitivity of 333.3 nm/RIU and a figure of merit (FOM) of 16.665 RIU^{-1} , aimed at the detection of various biomaterials (Farmani et al. 2020). Hamza et al.'s extensive investigations culminated in the innovation of a three-band microscale biosensor that employs Terahertz MIMs, demonstrating high sensitivity in identifying non-melanoma skin cancer (NMSC) (Hamza et al. 2024). Given the increasing prevalence of life-threatening conditions like cancer, it is imperative for researchers to broaden the application of optical biosensors for the benefit of society. Consequently, further theoretical and experimental investigations are essential for the advancement of optical biosensor design.

In recent years, there has been significant interest and exploration in the study of optical sensors utilizing split ring resonators (SRRs). This burgeoning field has seen the research and development of numerous SRR-based sensors crafted from plasmonic metamaterials, reflecting the growing recognition of their potential in various applications (Jafrasteh et al. 2023; Salehnezhad et al. 2023; Farmani et al. 2020; Hamza et al. 2024; Wellenzohn and Brandl 2015; Abdolrazzaghi et al. 2018; Horestani et al. 2020; Chen et al. 2020; Yang and Lin 2021). Amazing electromagnetic properties, like negative refractive index (Cao et al. 2020), the superlens effect (Chen et al. 2018; Song, et al. 2021) and invisible coating (Jing et al. 2020; Ye et al. 2021), are possessed by metamaterials. The characteristics of metamaterials are frequently interconnected with localized surface plasmon resonances (LSPR), which arise from the collective movements of unbound electrons generating intense electromagnetic fields near the metallic components of the metamaterial (Verma et al. 2024). Pendry et al. (1999) introduced the idea of metamaterial for the first time in 1999. Gold is a plasmonic metamaterial that is crucial to the operation of optical sensors based on SRR as a resonator (Song et al. 2019; Zaitsev et al. 2019). High-sensitivity biosensors are needed today to evaluate various biological samples. This is because there are many deadly diseases in the medical sciences, and diagnosing them calls for sensors with accurate measurement and increased sensitivity (Haq Khan et al. 2023). Throughout the years, a multitude of strategies have been investigated to improve sensitivity in plasmonic applications. These include surface plasmon resonance, optical fibers, optical waveguides, and meta surfaces (Kazanskiy et al. 2022). Because of their high sensitivity and label-free nature, refractive index sensors based on surface plasmon resonance (SPR) and surface plasmon polariton (SPP) currently have many applications in biological processes and medical sciences (Luo et al. 2014; Mandracchia et al. 2015; Wang and Fan 2016; An et al. 2017; Wang et al. 2022; Liu et al. 2023). But because SPR or SPP systems need optical couplers (like prisms and gratings), their production is labor-intensive and expensive due to the need for precise tools (Nootchanat et al. 2019; Benedikovic et al. 2019; Ropp et al. 2023; Laffont, et al. 2023; Flynn et al. 2023). In recent years, researchers have investigated different metal–insulator–metal configurations for plasmonic ring resonators. However, the presence of metal loss in these structures poses a significant challenge as it greatly impacts the quality factor (Wu et al. 2017). Vertical split-ring resonator-based nano plasmonic sensors have demonstrated significant advancements in greatly enhancing sensitivity (Wu et al. 2014). A circuit model for analyzing metal–insulator–metal plasmonic complementary split-ring resonators provides a comprehensive framework for understanding and optimizing their performance in various sensing applications (Bahadori et al. 2014). Optical slot-assisted meta surfaces have shown great potential for IgG protein detection, offering enhanced sensitivity and selectivity in biosensing applications (Brunetti et al. 2024). Here, we present a plasmonic sensor based on a split ring resonator (SRR) to greatly reduce the aforementioned limitations in order to address these issues.

Additionally, SRR structures have multiple reflection resonances, whereas most SPR or SPP structures are single-mode, meaning they cannot cover more than one resonance (Buzavaite-Verteliene et al. 2020; Li, et al. 2021). Applications for structures made with SRR-based metamaterials are numerous and span a wide range of industries. Although SRR structures are easier and less expensive to fabricate than SPR and SPP-based sensors, their sensing area is typically smaller and their FOM is lower. While numerous studies have been carried out to enhance the quality factor of SRR sensors, new types of SRR-based sensors have been presented to improve the sensing area (Ye et al. 2022). In (Ma et al. 2021), a three-stage coupled SRR structure was introduced, and in Mohammadi et al. (2021), an active feedback loop was studied as a means of improving sensor resolution. Nevertheless, these sensors had a comparatively low sensitivity. Applications for structures made with SRR-based metamaterials are numerous and span a wide range of industries. These include active filters, biosensors with high sensitivity for detecting diseases like cancer, and narrowband filters for wireless communication systems. The obtained results demonstrate that the SRR-based sensor has a relatively good sensitivity, FOM, and high-quality factor in the first mode, and high sensitivity and relatively good quality factor in the second mode. These characteristics make the sensor useful for understanding biological processes, diagnosing a variety of diseases, including cancer and in the pharmaceutical industry.

2 Structure and method

This section presents the design of the split ring (SRR) resonator-based multi-mode refractive index (MMRI) plasmonic biosensor as well as the simulation approach used to solve the governing equations. This allows for the comparison of performance metrics and the detection of distinct cancer cells in each mode. Figure 1a depicts the three-dimensional cell structure of the suggested sensor unit, which is made up of two split ring resonator (SRR) that are concentric on a quartz substrate, from both the top view (xy plane) as shown in Fig. 1b and the left view (xz plane) as shown in Fig. 1c. As can be seen, the outer SRR (SSR1) has side length d and the inner SRR (SRR2) has the outer radius r_{\max} and the inner radius r_{\min} and the width $W_r = r_{\max} - r_{\min}$. SRR2 is positioned within the air cylinder so that SRR1's center has a radius of R . SRR1 is a cube-shaped piece of gold with an air gap of g_1 and SRR2 is a cylinder-shaped piece of silver with an air gap of g_2 . h_a is the definition of the thickness of both SRR1 and SRR2. SRR2 will be separated from SRR1 by a distance of h_a , while SRR1 is positioned precisely on the quartz substrate. In the next stages of the work, the value of h and other structural dimensional parameters are scanned for the best possible performance of the suggested sensor. Additionally, the refractive index of the materials that were used—such as gold, silver, and quartz—was taken from the data of Coode Adams (Coode-Adams 1927), Johnson and Christy (Johnson and Christy 1972), and Palik (Handbook of optical constants of solids 1998), respectively. The effect of structural parameters refers to how variations in the geometric characteristics of the sensor device, such as dimensions, shape, or material composition, influence its performance and capabilities. In the context of a split ring resonator (SRR)-based multi-mode refractive index plasmonic biosensor, structural parameters may include the size and shape of the SRRs, the spacing between them, the material properties of the resonator components, and any other relevant geometric features. Table 1 displays the initial values for the structure's dimensional parameters.

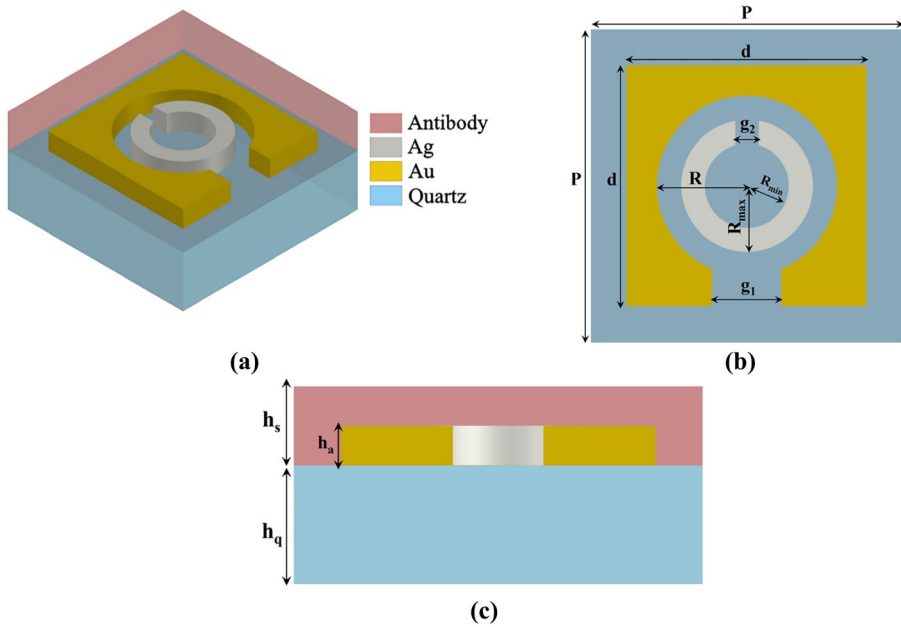


Fig. 1 **a** 3D schematic of the proposed structure, **b** Top view cross section and **c** Left side view cross section

Table 1 Dimensional parameters of the proposed structure

Parameter	Symbol	Value (nm)
Quartz side length	P	520
The length of the side of the outer disc	d	400
The length of the first air gap	g ₁	115
The length of the second air gap	g ₂	38.4
The radius of the air circle inside the disc	R	150
The outer radius of the ring	r _{max}	110
The inner radius of the ring	r _{min}	70
Inner ring width (r _{max} – r _{min})	W _r	40
Quartz height	h _q	150
Disc and ring height	h _a	50
The height of the ring of quartz	h	0–60
antibody height	h _s	100

To compare the performance parameters in each of the modes, the desired structure was simulated using the finite difference time domain (FDTD) numerical solution method. From the top view, a plane wave light source in various wavelengths is projected onto the xy plane with a polarization angle of 0 degrees. Along the x and y axes, periodic boundary conditions are chosen, and Perfectly Matched Layer (PML) boundary conditions are chosen along the z axis. The refractive index of the desired natural

Table 2 Refractive index values of different cancer cells

Cells	Cancer cells	RI (n)	References
Normal cell	-	1.350	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
Normal cell	Skin cancer	1.360	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
Basal		1.380	
Normal cell	Blood cancer	1.376	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
Jurkat		1.390	
Normal cell	Cervical cancer	1.368	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
Hela		1.392	
Normal cell	Adrenal gland cancer	1.381	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
PC12		1.395	
Normal cell	Breast cancer	1.385	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
MDA-MB-231		1.399	
Normal cell	Breast cancer	1.387	Sharma et al. (2015); Kumar et al. (2017); Jabin et al. (2019); Ahmed et al. (2021)
MCF-7		1.401	

antibody and different cancer cells are also shown in Table 2. Because the desired structure is created and simulated to detect various cancer cells.

The presence of cancer cells alters the normal antibody's refractive index (Δn_{s_s}), and this alteration in refractive index causes a shift in the structure's reflection spectrum's wavelength ($\Delta\lambda_0$). Consequently, the suggested biosensor's sensitivity (S) is defined as Eq. 1 (Bijalwan et al. 2021).

$$S = \frac{\Delta\lambda_0}{\Delta n_{s_s}} \tag{1}$$

Furthermore, the FOM and quality factor (Q-factor) of the biosensor are expressed as Eq. 2 and Eq. 3, respectively (Yang and Lin 2021; Jalil et al. 2021).

$$FOM = \frac{S}{FWHM} \tag{2}$$

$$Q\text{-factor} = \frac{\lambda_0}{FWHM} \tag{3}$$

where FWHM is the full width at half maximum.

3 Results and discussion

The proposed split ring resonator configurations are investigated to determine the optimal mode concerning material composition and dimensional attributes for the detection of diverse cancer cells. The determination of the most favorable resonant frequencies was conducted through simulations of the biosensor's reflection spectrum across varied wavelengths, as illustrated in Fig. 2, employing gold and silver as materials for SRR1 and SRR2. Examination of the results reveals two distinct scenarios where the most suitable resonances manifest: firstly, when both SRRs are fabricated from silver, and secondly, when SRR1 is composed of gold while SRR2 is made of silver. Figure 2a, e show the reflectance

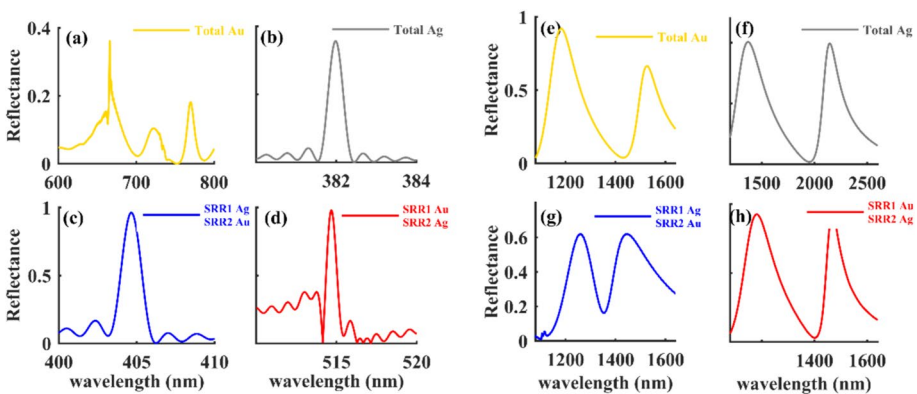


Fig. 2 a, e The reflection spectrum of the proposed structure in the first mode (visible wavelength) and the second mode (NIR wavelength) for both Au SRRs, b, f for both Ag SRRs, c, g for SRR1 made of Ag and SRR2 made of Au, and d, h for SRR1 made of Au and SRR2 made of Ag

spectra when both SRRs are made from gold in the visible wavelength and Near-infrared (NIR) wavelength ranges, respectively. Figure 2b, f show the same results when both are made from silver, while Fig. 2c, g illustrate the spectra for SRR1 made of silver and SRR2 made of gold. Finally, Fig. 2d, h present the spectra for SRR1 made of gold and SRR2 made of silver.

The first mode is chosen for the wavelength range of 300–800 nm, while the second mode is selected for the wavelength range of 1000–3000 nm to avoid redundancy. Figure 3a, b depict the contour plot of the reflection spectrum of the proposed structure within the visible and NIR wavelengths, taking into account the distance (h) of SRR2 from the quartz substrate, ranging from 0 to 60 nm. The analysis presented in Fig. 3b reveals that the resonances observed in the second mode exhibit relatively similar characteristics. On the other hand, the resonances in the first mode exhibit significant variations with changes in h , leading to alterations in the resonance wavelength range. The influence of structural parameters on multimodal resonances is a complex interplay involving factors such as geometrical dimensions, material properties, and the electromagnetic environment. Changes in structural parameters like the dimensions of the split ring resonator (SRR), the distance between the SRRs, and the composition of the materials can significantly impact the resonant behavior of the system. For instance, altering the dimensions of the SRRs can lead to shifts in resonant frequencies due to changes in the capacitance and inductance of the structure, affecting the coupling between different modes. Similarly, modifying the distance between SRRs can influence the electromagnetic coupling between them, thereby affecting the mode distribution and resonance frequencies. Furthermore, varying the material composition can change intrinsic properties such as conductivity and permittivity, consequently altering the electromagnetic response of the structure. Understanding the intricate relationships between these parameters is crucial for optimizing multimodal resonances in applications such as biosensors, where precise control over resonance characteristics is essential for sensitive detection.

The sensitivity analysis revealed that changes in SRR dimensions had a significant impact on resonant wavelength. Furthermore, variations in material properties were found to alter the electromagnetic response of the structure, resulting in shifts in resonance frequencies and changes in mode distribution. The following examines how variations in SRR2's width (W_r), antibody thickness (h_s), and distance (h) from the quartz substrate

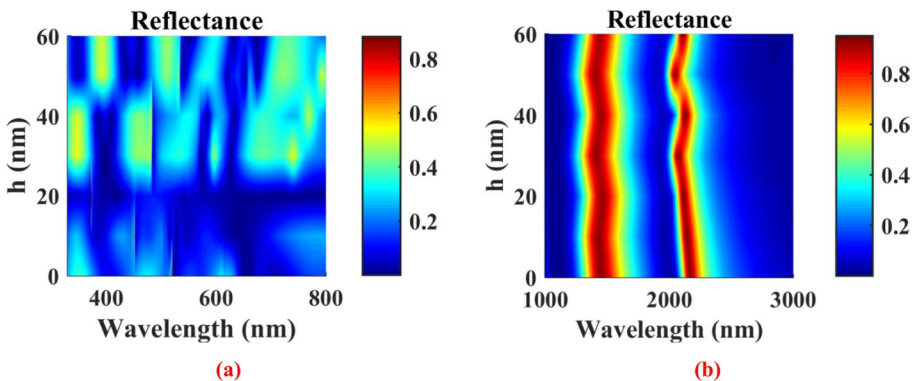


Fig. 3 **a** contour plot of reflected responses as function of wavelength and h from 0 to 60 nm in the visible wavelength and **b** in the NIR wavelength

affect the sensor's sensitivity and figure of merit for the refractive index of the antibody from 1.33 to 1.41 in one of the resonance modes, from 1000 to 3000 nm. The resonance wavelength diagram in Fig. 4a–c is drawn for $W_r=20\text{--}60$ nm, $h=0\text{--}60$ nm, and $h_s=80$ nm–180 nm, respectively. It represents the antibody refractive index from 1.33 to 1.41 in one of the resonance modes, from wavelength 1000 nm to 3000 nm. The sensitivity of the intended sensor is indicated by the graphs' slope. The diagram in Fig. 4a shows that $h=60$ nm corresponds to the line with the largest slope based on observations. Additionally, the line with the largest slope in Fig. 4b corresponds to $W_r=20$ nm, whereas the line in Fig. 4c does not differ significantly when h_s changes from 80 to 180 nm.

The figure of merit serves as a metric for optimization efforts. Researchers can use it to identify parameters or design features that have the most significant impact on performance and focus their efforts on optimizing those aspects. The sensitivity and FOM diagram for h_a changes from 0 to 60 nm are shown in Fig. 5a. As you can see, our preferred wavelength of $h=60$ nm had the highest sensitivity and FOM. The impact of variations in SRR2's width (W_r) on the sensor's sensitivity and FOM is examined in the following stage. We only alter r_{\min} in this investigation; r_{\max} remains constant at 110 nm. Figure 5b from this investigation demonstrates that at $W_r=20$ nm, the greatest sensitivity and the smallest FOM occur, and at $W_r=60$ nm, the smallest sensitivity and the largest FOM occur. In order to maximize FOM and sensitivity, the best W_r , which is 40 nm, is chosen.

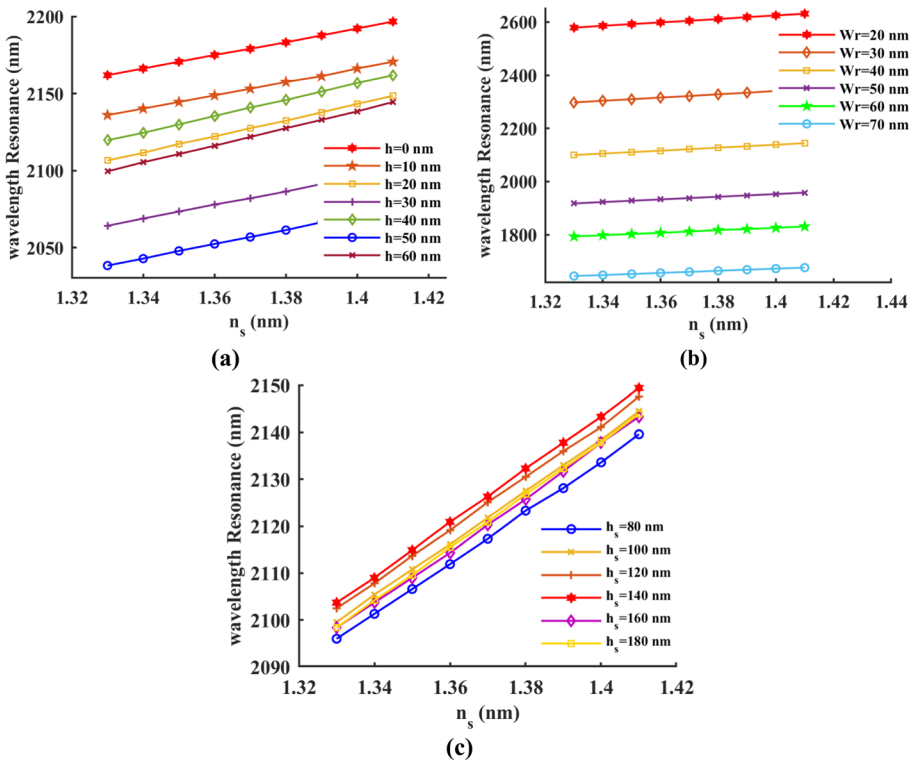


Fig. 4 **a** Resonance wavelength diagram for antibody refractive index in the second state for h from 0 to 60 nm, **b** for W_r from 20 to 70 nm, and **c** for h_s from 80 to 180 nm

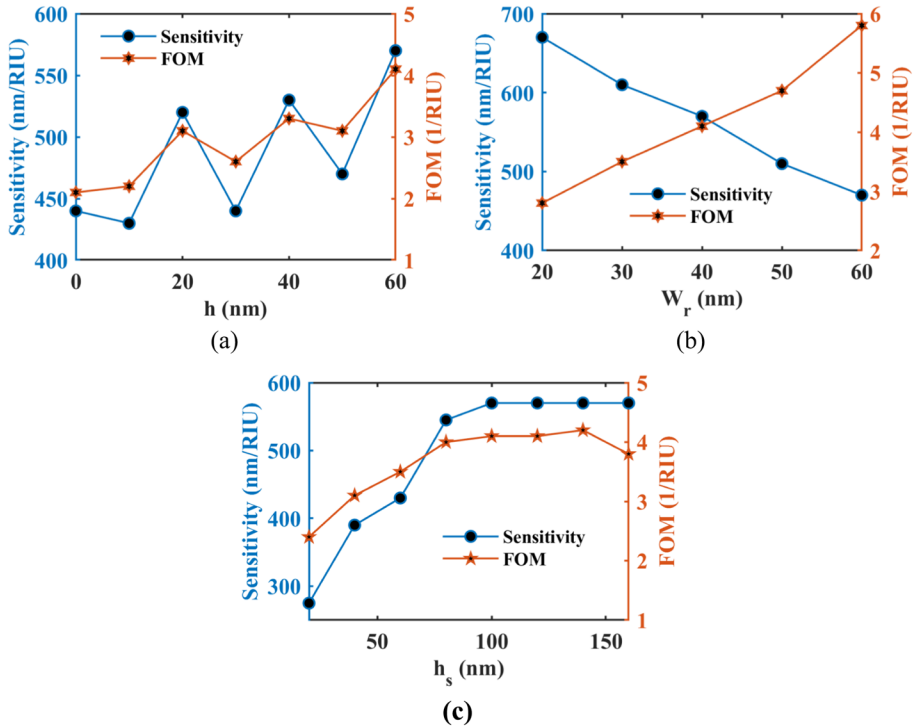


Fig. 5 **a** sensitivity and FOM in the second mode for h from 0 to 60 nm, **b** per W_r from 20 to 60 nm, and **c** per h_s from 20 to 160 nm

This step looks into how the sensitivity and FOM of the sensor are affected by variations in the desired antibody's (h_s) thickness. The corresponding diagram, shown in Fig. 5c, shows that, in accordance with the results, the sensor's sensitivity and FOM increase from $h_s=20$ nm to $h_s=100$ nm. However, once h_s exceeds 100 nm, the sensor's sensitivity and FOM virtually remain constant. As a result, we set the antibody's thickness (h_s) to 100 nm in this instance. It should be noted that for the investigations conducted in each of these steps, we utilized the second mode described above.

We first evaluate the biosensor in one of the first modes, ranging in wavelength from 300 to 800 nm, in order to identify various cancer cells, after determining the right dimensions for the constructed structure. The reflection spectrum of this state is shown in Fig. 6a. Table 3 displays the outcomes from this section as well. This section yielded the highest sensitivity and FOM, which are 82.5 nm/RIU and 20.53 1/RIU, respectively. These results are superior to those of the biosensor suggested in Ye et al. (2022), which is based on crystal photonics.

Next, the performance parameters are analyzed in second mode, in wavelength ranging from 1000 to 3000 nm. Figure 6b displays the reflection spectrum of the aforementioned state, and Table 4 reports the obtained results. This section shows that the computed FOM and maximum sensitivity are equal to 4.1 1/RIU and 570.4 nm/RIU, respectively. Additionally, it has more optimal results for prostate cancer (PSA) detection than the structure suggested in Shokorlou et al. (2022), which is based on metal–insulator–metal rings.

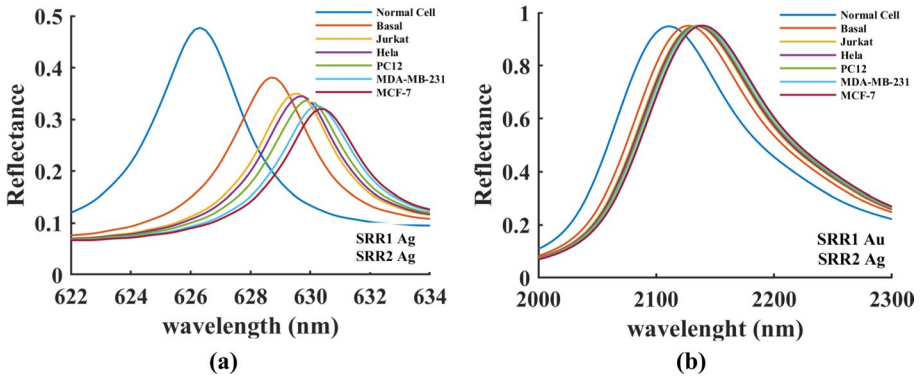


Fig. 6 **a** The reflection spectrum of the proposed biosensor for the refractive index of cancer cells in the first mode and **b** the second mode

Table 3 Performance parameters of the proposed biosensor in the first mode

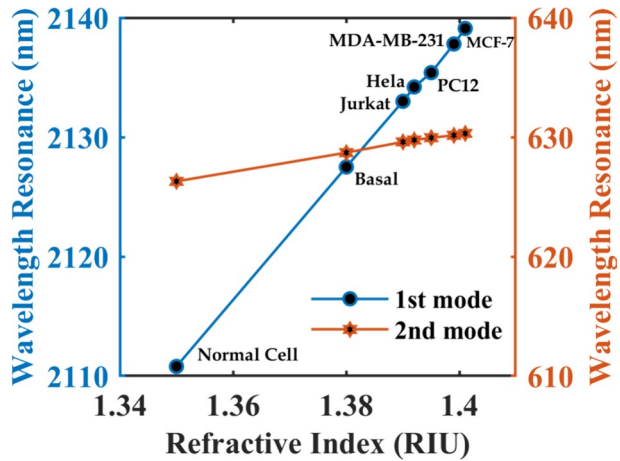
Cells	Cancer cells	RI (n)	Sensitivity (nm/RIU)	FWHM (nm)	λ_0 (nm)	Q-factor	FOM
Normal cell	–	1.350	–	–	626.30	–	–
Normal cell	Skin cancer	1.360	75.0	4	627.20	157.17	18.75
Basal		1.380			628.70		
Normal cell	Blood cancer	1.376	78.5	4.18	628.50	150.60	18.78
Jurkat		1.390			629.60		
Normal cell	Cervical cancer	1.368	82.5	4	627.78	157.44	20.53
HeLa		1.392			629.76		
Normal cell	Adrenal gland cancer	1.381	75.0	4.3	628.90	146.26	17.45
PC12		1.395			629.95		
Normal cell	Breast cancer	1.385	79.3	4.15	629.04	151.58	19.10
MDA-MB-231		1.399			630.15		
Normal cell	Breast cancer	1.387	77.9	4.18	629.22	150.53	18.63
MCF-7		1.401			630.31		

Furthermore, compared to values reported in works (Panda et al. 2021; Arafa et al. 2017), the sensitivity of the proposed sensor is significantly higher. When comparing the data from the first and second cases, we find that the sensor’s sensitivity in the second case is significantly higher than in the first, but this is not the case for FOM. The resonance wavelength diagram for the cancer cells’ refractive index in the two cases under examination is also displayed in Fig. 7. The calculated sensitivity indicator is represented by the slopes of the lines drawn in this diagram; as you can see, the second case’s line has a much higher slope than the first case’s, which is consistent with the earlier explanation. So, we optimized the design of the plasmonic split ring resonator biosensor by varying key structural parameters such as the ring dimensions, gap size, and material properties. These modifications directly influence the resonant modes of the sensor. Specifically, changes in these parameters affect the electromagnetic field distribution and coupling effects within the resonator, which in turn alter the resonance characteristics. For example, adjusting the gap size can change the strength of the electric field in the split region, thereby shifting the

Table 4 Performance parameters of the proposed biosensor in the second mode

Cells	Cancer cells	RI (n)	Sensitivity (nm/RIU)	FWHM (nm)	λ_0 (nm)	Q-factor	FOM
Normal cell	–	1.350	–	–	2110.8	–	–
Normal cell	Skin cancer	1.360	570.40	139.22	2116.1	15.28	4.1
Basal		1.380			2127.5		
Normal cell	Blood cancer	1.376	561.30	141.2	2125.1	15.1	4
Jurkat		1.390			2133.0		
Normal cell	Cervical cancer	1.368	553.35	140.55	2120.9	15.18	3.95
Hela		1.392			2134.2		
Normal cell	Adrenal gland cancer	1.381	519.50	142.30	2128.1	15	3.65
PC12		1.395			2135.4		
Normal cell	Breast cancer	1.385	520.70	142.50	2130.6	15	3.66
MDA-MB-231		1.399			2137.8		
Normal cell	Breast cancer	1.387	564.50	142.80	2131.2	14.98	3.95
MCF-7		1.401			2139.1		

Fig. 7 Resonance wavelength for refractive index of cancer cells in the first mode and the second mode



resonance frequency and enhancing sensitivity to refractive index changes. Similarly, varying the ring dimensions affects the effective optical path length and can lead to changes in the quality factor and figure of merit. These optimizations are crucial as they directly impact the sensor’s performance in terms of sensitivity for different cancer cell types. By systematically studying and fine-tuning these parameters, we have achieved a design that is a multi-modal biosensor.

Two resonances have been found in the reflection spectrum of Fig. 3b, corresponding to the second state. To better understand the electromagnetic responses of the biosensor at these wavelengths, we examine the distribution of the electric and magnetic fields of both SRRs under the assumption that $h=0$ nm. The distribution of the electric and magnetic fields at wavelengths of 1367 nm and 2143 nm are depicted in Fig. 8a, b, both in the xy-cross section and from the top view. The magnetic field is primarily dispersed in the area between the two SRRs, while the electric field is primarily distributed inside the gap between SRR1 and SRR2, as well as along the outer walls of SRR1. Additionally, we show

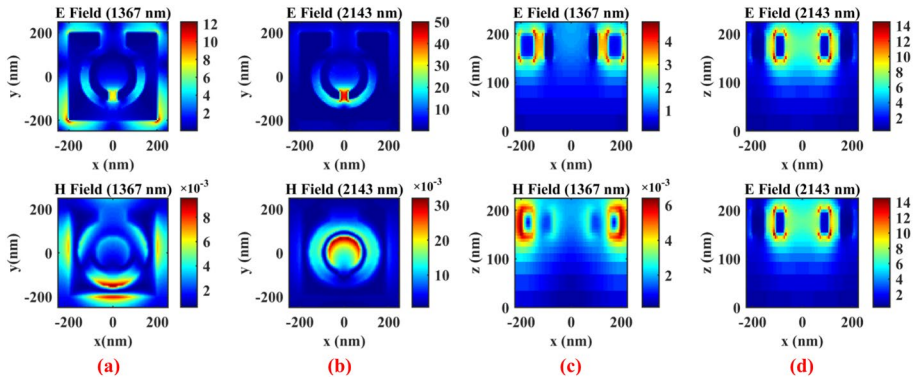


Fig. 8 **a, c** Electric and magnetic field distribution of the sensor for the xy and xz planes, respectively, at the wavelength of 1367 nm. **b, d** wavelength 2143 nm

the distribution of the electric and magnetic fields in the xz-cross section and from the left view at wavelengths of 2143 nm and 1367 nm in Fig. 8c, d. To optimize the design of the plasmonic split ring resonator biosensor, we varied key structural parameters such as the ring dimensions, gap size, and material properties. These modifications directly influence the resonant modes of the sensor, which, in this case, are Lorentzian resonant modes.

Figure 9 illustrates the reflection, transmission, and absorption characteristics for both of visible and NIR wavelength ranges. Specifically, Fig. 9a shows the reflection, transmission, and absorption diagram for the wavelength range of 300–800 nm, while Fig. 9b covers the wavelength range of 1000–3000 nm. According to these spectra in visible and NIR wavelength ranges, the proposed structure can be used in optical applications, including filters, absorbers and imaging.

The sensor that was designed can be used to detect a variety of diseases, including cancer, as well as environmental processes. It has a high sensitivity in the second mode and a relatively good sensitivity and high FOM. The sensitivity and FOM values can be significantly increased by increasing the number of internal SRRs in the suggested structure (Yang and Lin 2021). Also, Eq. 3, which received less attention in earlier published works,

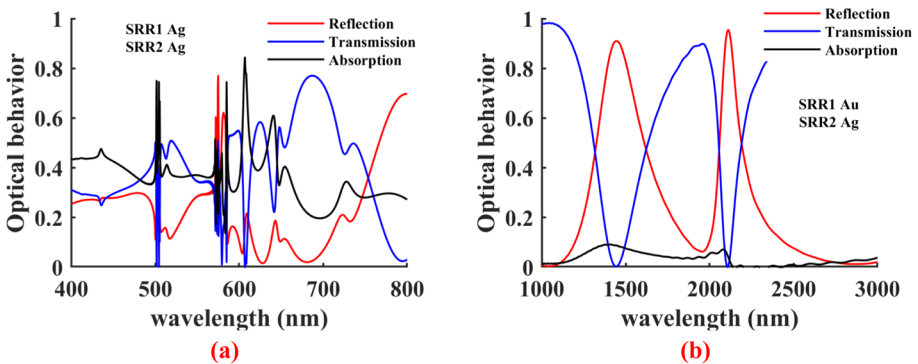


Fig. 9 **a** Reflection, transmission and absorption diagram for the wavelength range of 400–800 nm, **b** For the wavelength range of 1000–3000 nm

was used in this study to examine the quality coefficient of the designed sensor. Based on the data presented in Tables 3 and 4, the first case's quality factor is significantly higher than the second case's, where the maximum quality factor of 155.44 was achieved. So, our proposed plasmonic SRR biosensor exhibits a higher sensitivity in second mode and higher figure of merit in first mode, which compares favorably with other recently reported sensors. The comparison of sensor performance parameters in both modes with previous works is reported in Table 5. By addressing the current limitations of plasmonic biosensors and leveraging the advantages of SRR design, our proposed sensor provides a balanced solution with improved sensitivity, manufacturability, and practical applicability, making it a strong candidate for early cancer detection and other biomedical applications.

Several approaches exist for the possible development and fabrication of the suggested structure. Nanofabrication methods like electron beam nanolithography (EBL), focused ion beam lithography (FBL), and nanoimprint lithography are among the potential techniques that can be utilized. These techniques offer the opportunity to attain accurate patterning of different materials essential for creating Subwavelength Resonators (SRRs). The sequential assembly of materials necessary for the construction of SRRs can be accomplished by examining various techniques, including layer-by-layer deposition, self-assembly, and transfer printing. These approaches allow for the transfer of prefabricated SRRs from an initial substrate to the final sensor substrate, ensuring smooth integration with various materials. This integration improves the compatibility and performance of the SRRs in the sensor.

4 Conclusion

In this study, we developed and simulated a multimode refractive index (MMRI) plasmonic biosensor utilizing a split ring resonator (SRR) architecture for the detection of various cancer cells. Leveraging the Finite-Difference Time-Domain (FDTD) numerical solution method, we analyzed the designed biosensor. First SRR is a cube-shaped piece of Au with an air gap and second SRR is a cylinder-shaped piece of Ag with an air gap. The effect of structural parameters such as dimensions, shape, or material composition, on performance parameters of proposed structure were investigated. Demonstrating promising

Table 5 Comparison of the performance parameters of the proposed structure with previous works

References	Base	Sensitivity (nm/RIU)	FWHM (nm)	λ_0 (nm)	Q-factor	FOM (1/RIU)
Shokorlou et al. (2022)	MIM	567.23	152.35	1530	–	3.72
Panda et al. (2021)	PC	290	–	621.5	–	1074.04
Arafa et al. (2017)	PC	462	–	1600	$1.11 \cdot 10^5$	–
Farmani et al. (2020)	SPR	333.3	20	1200	–	16.665
Bijalwan et al. (2021)	PC	74.5	–	825	–	19.6
Shi et al. (2022)	SMRR	403	–	1560	–	–
Goede et al. (2019)	MRR	100	–	1565	$3.2 \cdot 10^5$	–
Jian et al. (2017)	MRR	200	–	1560	10^4	–
This work (1st mode)	SRR	82.5	4	629.76	157.44	20.53
This work (2nd mode)	SRR	570.4	139.22	2127.5	15.28	4.1

performance, the biosensor exhibited a sensitivity exceeding 80 nm/RIU and a Figure of Merit (FOM) surpassing 20 RIU⁻¹ in the first mode, while in the second mode, it displayed a sensitivity exceeding 570 nm/RIU and an FOM exceeding 4 RIU⁻¹. These results highlight the dual benefits of the biosensor: the first mode provides higher precision and accuracy due to its superior FOM, making it ideal for applications requiring detailed and reliable measurements. Meanwhile, the second mode's high sensitivity enables the detection of minute changes in refractive index, which is crucial for early-stage cancer detection and other medical diagnostics. This dual-mode capability enhances the versatility and effectiveness of the biosensor, potentially leading to significant advancements in the field of cancer diagnostics and treatment. The ability to detect and measure small refractive index changes with high sensitivity and precision underscores the biosensor's potential for broad applications in biomedical research and clinical settings.

Author contributions All authors contribute equally.

Funding The authors have not disclosed any funding.

Data availability This manuscript does not report data generation or analysis.

Declarations

Conflict of interest The authors declare no competing interests.

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