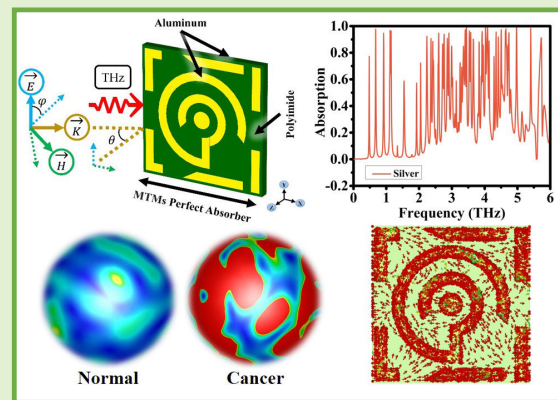


Development of a Terahertz Metamaterial Micro-Biosensor for Ultrasensitive Multispectral Detection of Early Stage Cervical Cancer

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Abstract—This research introduces an innovative design for a metamaterial (MTM)-based compact multiband biosensor aimed at early stage cervical cancer detection. The device operates within the terahertz (THz) frequency range, specifically from 0 to 6 THz. The proposed sensor architecture features an MTM layer composed of a patterned aluminum structure deposited on a polyimide substrate. The primary design objective is to optimize the geometry parameters to achieve near-perfect absorption of electromagnetic (EM) waves across the entire operating bandwidth. The design process utilizes full-wave EM simulation tools. This article details all intermediate steps in the sensor's topology development, guided by an investigation of the absorption characteristics of successive architectural variations. It also analyzes the effects of the substrate and resonator material. The suitability of the proposed sensor for early stage cancer diagnosis is demonstrated using a microwave imaging (MWI) system that incorporates the device. Extensive simulation studies confirm the sensor's capability to distinguish between healthy and cancerous cervical tissue. For further validation, comprehensive benchmarking is conducted against numerous state-of-the-art sensor designs reported in recent literature. These comparative studies indicate that the proposed sensor offers superior performance in terms of absorbance levels and the width of the operating bandwidth, both of which enhance the sensitivity of cancer detection.

Index Terms—Biosensor, cervical cancer, metamaterials (MTMs), microwave imaging (MWI), multiband sensors.



I. INTRODUCTION

TERAHERTZ (THz) waves possess a range of unique advantages that make them highly promising for various applications, particularly in biological detection and pharmaceutical fields. The combination of low photon energy and

strong penetration ability akin to X-rays makes THz waves particularly well suited for exploring biological tissues and molecules. One of the key advantages of THz technology lies in its ability to perform label-free, nondestructive, and noncontact detection, thus preserving the integrity of analyte for further analysis [1], [2]. Furthermore, THz sensors offer real-time monitoring capabilities, enabling researchers to observe dynamic biological processes as they unfold [3]. This real-time feedback is invaluable in pharmacological applications, where rapid assessment of drug interactions and effects is critical for developing effective treatments. Moreover, the nontoxic and nondestructive nature of THz waves to human tissue makes them safer for use in medical diagnostics compared to other techniques that may involve ionizing radiation or invasive procedures, thus minimizing potential health risks associated with diagnostic procedures [4]. Overall, the unique properties of THz technology, including its noninvasive

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nature, real-time monitoring capabilities, and ability to provide detailed information about biological samples, position it as a highly promising tool for a wide range of applications in biomedical research, pharmaceutical development, and clinical diagnostics.

Metamaterials (MTMs) represent a cutting-edge class of artificial electromagnetic (EM) materials constructed from subwavelength units arranged in a periodic array. These materials are engineered to exhibit unique EM properties not found in naturally occurring substances such as negative refractive index and negative permittivity. These properties enable MTMs to manipulate EM waves in unconventional ways, leading to applications such as absorbers [5], active filters [6], and ultrasensitive biosensors [7], [8], [9], [10]. In addition, MTMs can exhibit inverse Doppler effects [11], which opens up new possibilities in signal processing and communication systems. Resonant MTMs, in particular, possess micro-sized gaps that enable them to concentrate and enhance the intensity of local EM fields. This enhancement is advantageous for applications requiring high sensitivity to changes in dielectric properties, making resonant MTMs particularly well suited for THz sensing. In THz sensing, the ability to detect minute changes in dielectric properties is crucial for identifying and analyzing biomolecules, chemical substances, and other materials [12]. By leveraging the enhanced sensitivity provided by resonant MTMs, THz sensors can achieve higher accuracy and resolution in detecting and characterizing samples [13]. Overall, the unique EM properties of MTMs, particularly resonant MTMs, make them powerful tools for advancing THz sensing technologies and opening up new possibilities in fields such as biomedical diagnostics specifically cancer cell sensing [14], [15] and chemical sensing [16], [17], [18], [19], [20], [21], [22], environmental monitoring, materials science, and communication systems.

With millions of new cases diagnosed each year and a substantial number of deaths attributed to the disease, cancer remains a major public health concern worldwide and also a foremost inhibitor to increasing life expectancy. The World Health Organization (WHO) report for global cancer data released in 2020 indicated an estimated number of 19.29 million new cancer cases worldwide, including 9.96 million deaths. It is expected this number of new cases to increase to 28.4 million in 2040 [23], [24]. Efforts to address cancer incidence and mortality include not only advancements in treatment methods but also initiatives focused on prevention, early detection, and improving access to healthcare services. The existing techniques, such as electrochemical methods, histopathological or immunological methods, fluorescence imaging, and cytometry, suffer from limitations such as lack of specificity, high costs, lengthy operation times, and the need for skilled personnel [25], [26]. However, surface plasmon resonance-based MTM sensors indeed offer several advantages over traditional methods for cancer cell detection. Their high precision, label-free detection, real-time measurement capabilities, and minimal sample requirement make them promising candidates for improving cancer diagnostics. Recently, a number of research works reported MTM-based sensors in THz range for cancer cell detection [14], [15], [27], [28], [29],

TABLE I
COMPLETE LIST OF THE VARIABLES THAT HAVE BEEN ADJUSTED FOR THE RECOMMENDED SENSOR

Parameter	Value (μm)	Parameter	Value (μm)
A	49	J	11.6
B	58	K	10
C	10.25	L	55
D	39.3	M	150
E	21.8	N	150
F	20.5	Aluminum (Al) thick	0.2
G	49	Polyimide thick (T1)	10
H	75.8	Coverslip thick (T2)	4
I	30	HeLa Cells thick (T3)	8

[30], [31], [32], [33], [34], [35], [36], [37], [38], in addition to sensors for various other biodetection applications [16], [17], [18], [19], [20], [21], [22], [39], [40], [41], [42], [43], [44], [45], [46], [47]. These reported biosensors for cancer detection are mostly graphene-based [14], [27], [29], [30], while water-based THz MTM absorbers [15] are also presented. Likewise, several ring-based resonator structures [27], [33], [35], [36], [38] for MTM THz sensors are proposed for cancer cell detection.

This research presents a novel MTM-based compact multi-band biosensor for early stage cervical cancer detection, operating from 0 to 6 THz. The sensor features a patterned aluminum structure on a polyimide substrate, optimized for near-perfect absorption. Using full-wave EM simulations, the design process examines absorption characteristics and material effects. The sensor's efficacy is demonstrated with a microwave imaging (MWI) system, distinguishing between healthy and cancerous tissue. Extensive benchmarking against state-of-the-art sensors shows superior performance in absorbance levels and bandwidth, enhancing cancer detection sensitivity.

II. LAYOUT OF THE MODEL'S UNIT CELL AND INCIDENT FIELD CONFIGURATION

The effective conversion of applied energy into absorption is possible due to the unique MTM-inspired structure, as shown in Fig. 1. Perfect absorbers effectively decrease the reflection of incoming energy by absorbing the majority of it. In addition, the design of the MTM minimizes destructive interference, which further enhances absorption and weakens transmitted waves. The absorber is built using a commercial full-wave finite integration technique (FIT) that relies on a high-frequency EM solver. This allows for the study of MTMs across different frequency ranges and limits. Scientists have used numerical analysis to investigate structures such as perfect electric and magnetic conductors (PEC and PMC), unit cells, periodic arrays, and empty space. Fig. 1(a) shows a first resonator and Fig. 1(b) shows a second resonator, respectively. The proposed design shown in Fig. 1(c) represents a creation based upon a combination of first and second resonators. The overall size of the structure is $150 \times 150 \mu\text{m}$. All dimensions of the proposed structure are represented in Table I.

Fig. 2 illustrates the structural configuration and field directions for an MTM perfect absorber designed to function within the THz frequency range of 0–6 THz. Addressing

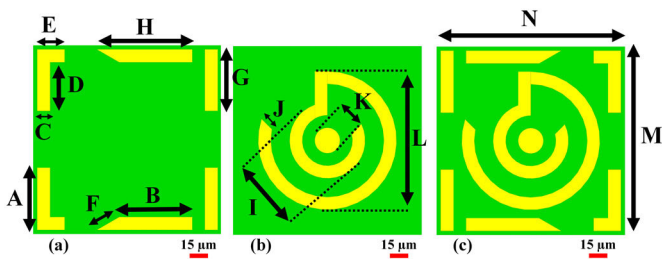


Fig. 1. Suggested structural design for a perfect absorber. (a) First resonators, (b) second resonators, and (c) proposed design.

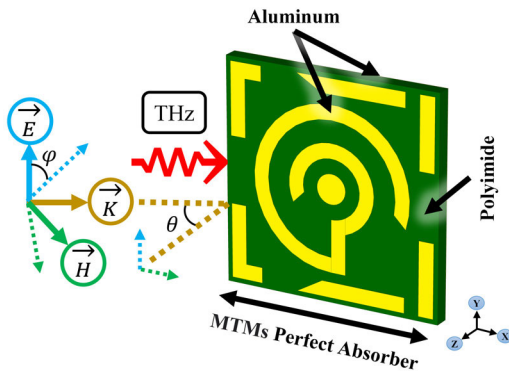


Fig. 2. Configuration of the structural design and the directions of the incident field for a perfect absorber.

the feasibility of high-frequency characterization, the study acknowledges the challenges associated with conventional electronic measurement techniques beyond 100 GHz. However, alternative optical methods, such as THz time-domain spectroscopy (THz-TDS), offer viable solutions for precise characterization within the 0.1 THz to tens of THz range. LTEM and thermal/pyroelectric techniques also present potential alternatives for intensity measurements [48], [49]. These methodologies while requiring further investigation, expand the possibilities for characterizing the proposed THz MTM biosensor. The design comprises an MTM layer, which includes a patterned aluminum structure deposited on a polyimide substrate. This configuration is optimized to achieve near-perfect absorption of incident EM waves over a wide bandwidth. The incident EM wave, depicted by the red arrow, interacts with the absorber surface at specific angles, denoted by θ and φ . The directions of the electric field (E), magnetic field (H), and wave propagation vector (k) are indicated by their respective arrows. These fields are essential for analyzing the absorber's performance, as the interaction between the incident wave and the MTM structure determines the absorption characteristics. The MTM perfect absorber effectively converts the incident EM energy into heat, minimizing reflection and transmission, thereby enhancing the sensing capabilities for applications such as biosensing and spectroscopy in the THz frequency domain.

III. RESULTS AND ANALYSIS

In this section, a comparative analysis is conducted covering compact absorber-based sensors tailored for THz operation aiming to obtain an optimized solution for cervical cancer

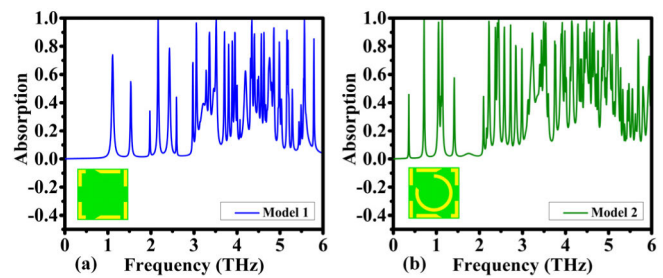


Fig. 3. Showing the different absorption characteristics of two designs. (a) Model 1. (b) Model 2.

diagnosis. To provide a thorough exploration of various architectural possibilities, this study progressively simulates, and assesses several distinct component designs. These designs are scrutinized based on their absorption spectrum, power distribution, and field density.

A. Absorption Characteristics of Different Designs

The first design under consideration was created from a modified square-shaped split-ring resonator, which is illustrated in Fig. 1(a) and it is sized around $150 \mu\text{m}$. To evaluate the performance of the structure, its absorption curve in function of the frequency was computed and the results are shown in Fig. 3(a).

This first model was already an operational design and showed interesting characteristics, as it had several absorption peaks with high absorption coefficients, including regions around 2, 3, 3.5, 4, and 5.5 THz.

A multiband absorber for these upper THz frequencies is of great interest for sensors in several systems, but there is still a desire to have absorption peaks in the frequencies around 1 THz as it presents different propagation behavior in the tissue. Due to that, a new set of designs was evaluated now using the obtained square resonator in conjunction with other structures, initiating with circular path-shaped structures and extending to a mix between circular split rings and bullseye patterned resonators.

The second developed model was evaluated and its results are provided in Fig. 3(b) together with an insert of its physical design. The absorption curve of this design shows great improvement in the frequencies around 1 THz, where at least three peaks clearly obtained very high absorption which partially solves the desired points, but some regions around 3 THz became worse and the very high frequencies over 5 THz are not being well represented too. Trying to achieve these specifications, a circular absorber according to the upper explanations was tried. Its physical constitution is shown in Fig. 1(b) and the resulting absorption spectrum is presented in Fig. 4(a).

The results of this third model were adequate just in the 3–5-THz range and did not show appropriate improvement in the lower or higher frequencies; in this way, the squared resonator was maintained and understood as an important part of the solution, especially for the higher part. In this way, a final model mixed Model 3 and the first model was developed.

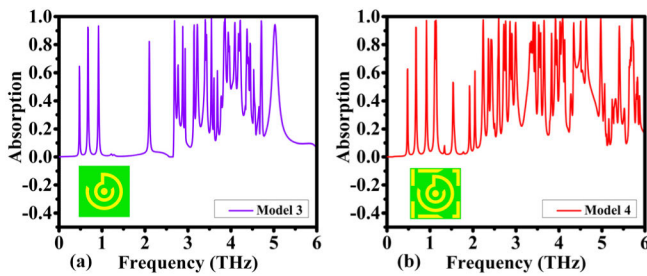


Fig. 4. Different absorption characteristics of the two designs. (a) Model 3. (b) Model 4.

The composition of this last model is shown in Fig. 1(c) and the resulting absorbance spectrum is presented in Fig. 4(b). This model achieved very good results, solving the absence of perfect absorbance peaks near 6 THz while also having several peaks with perfect absorption in the midband and covering the lower 1-THz region. In this way, this design was then chosen as the proposed model for this work and will be further evaluated with respect to its structural properties in the following subsections.

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The absorption intensity of the peaks is a deliberate feature intended to enhance the sensitivity of a biosensor. This heightened sensitivity is of paramount importance, particularly in practical applications, where accurate discrimination between healthy and cancerous cell samples is essential. The intensified peaks increase the opportunities for interaction with biomolecules, thereby facilitating a more definitive distinction between the two sample types. Multiple peaks correspond to the sensor's multiband functionality, allowing it to detect and differentiate between THz wavelengths for versatile sensing applications, as compared to single-band sensors offering limited applications. The multiband sensor can effectively absorb EM radiations at multiple frequencies, allowing for the operation across a wide frequency range with a comprehensive assessment of sensitivity across all bands. Thus, the multiband configuration enhances the sensitivity of the biosensor at discrete multiple frequency bands.

Although single-band models offer simplicity during fabrication and measurement, however, they suffer from limitations

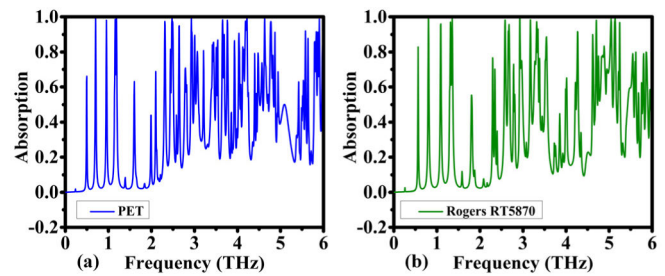


Fig. 5. Absorption spectra for the proposed design under different substrates. (a) PET. (b) Rogers RT 5870.

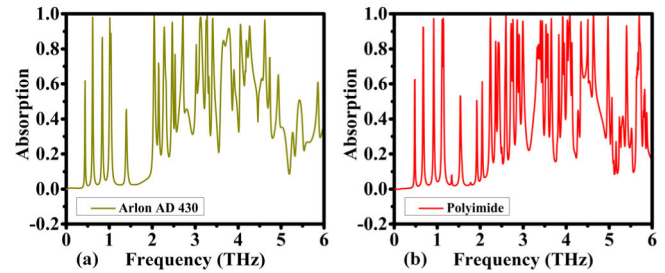


Fig. 6. Absorption spectra for the proposed design under different substrates. (a) Arlon AD 430. (b) Polyimide.

in sensitivity to diverse biomolecular interactions. The preference for the multiband configuration lies in its heightened sensitivity, specificity, and versatility. The multiband approach enables a more thorough and detailed analysis of THz interactions with biological samples, addressing the complexities of biomolecular signatures and minimizing cross-sensitivity, thus establishing it as the preferred choice for advanced biosensing applications.

B. Influence of Substrate and Resonator Material

MTM absorbance characteristics can be very sensitive to its composing materials and aiming to find an adequate material choice, the influence of substrate and resonator material was evaluated, trying to obtain an optimal balance of its ease of use, availability, and resulting characteristics.

The analysis started focusing on the substrate material, where PET and Rogers RT 5870 were evaluated for their absorption characteristics in Fig. 5(a) and (b), respectively. PET produced acceptable results maintaining several near-perfect absorption peaks. On the other hand, the RT 5870 did not work well in high frequencies. Other two materials were also evaluated, the Arlon AD 430 and polyimide in Fig. 6(a) and (b), respectively.

Arlon AD 430 performed even worse than the RT 5870, but polyimide was very successful, being slightly superior to PET, being easy to integrate, and providing good results in both low, middle and high, and high bands. The design was then evaluated with respect to its resonator material, where gold, silver, copper, and aluminum were simulated and its absorption results are presented in Figs. 7(a) and (b) and 8(a) and (b).

The influence of the resonator metal was found to be minor in relation to the changes eventuated in the substrate

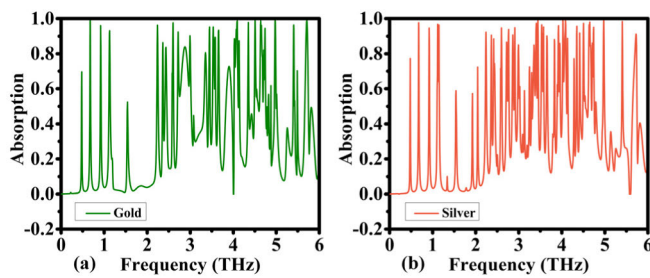


Fig. 7. Absorption spectra for the proposed design under various resonator material conditions. (a) Gold. (b) Silver.

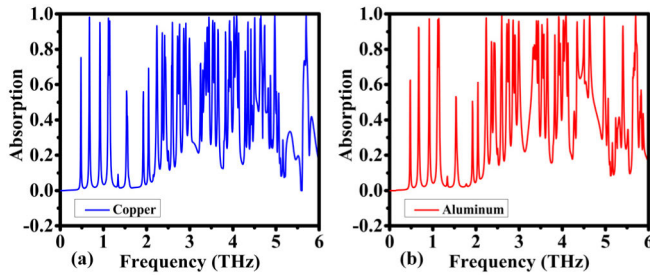


Fig. 8. Absorption spectra for the proposed design under various resonator material conditions. (a) Copper. (b) Aluminum.

material analysis. All the four metal options evaluated worked adequately having several peaks in all ranges well above 0.8. Gold and aluminum were the best metals, but the easiness of fabrication, low cost, higher availability, and better lithography and process integration favor aluminum, which was the final selected material together with the polyimide substrate to form the proposed micro-biosensor. The following subsections evaluate the substrate thickness selection.

C. Effect of Substrate Thickness

The performance of sensors depends on substrate thickness can affect their absorptivity. Research shows that variations in substrate thickness have a direct effect on the level of sensor absorptivity [50]. Sensors' sensitivity and detection capacities can be affected by thicker substrates' effects on their optical characteristics [51]. In addition, in some sensor applications, the thickness of the substrate influences horizontal magnetoresistance, which in turn affects the ability to detect horizontal magnetic fields (H -field) [52]. Substrate design, particularly thickness, is vital for attaining high sensitivity, stabilization, and homogeneity of molecules on the substrate in the context of carbon-based surface-enhanced Raman scattering (SERS) sensors. This has a substantial impact on the sensor's enhancement factor and detection capability [53].

A significant element that profoundly impacts the ability of sensors to absorb and function effectively across various applications is the thickness of the substrate. The investigation illustrated in Figs. 9–12 is focused on exploring the influence of substrate thickness on absorption capacity within a specific frequency range from 0 to 6 THz. Each Figure displays different substrate thicknesses and their respective responses, providing insight into how different thicknesses impact the

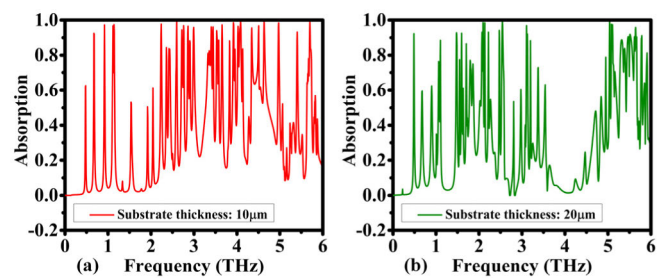


Fig. 9. Investigating the effect of the proposed substrate thickness on the ability to absorb (a) substrate thickness 10 μm and (b) substrate thickness 20 μm .

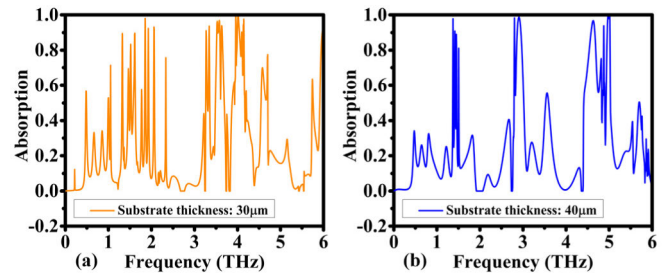


Fig. 10. Investigating the effect of the proposed substrate thickness on the ability to absorb (a) substrate thickness 30 μm and (b) substrate thickness 40 μm .

capacity to absorb EM waves. An initial observation from Fig. 9(a) and (b) with substrate thicknesses of 10 and 20 μm reveals a favorable response throughout the frequency spectrum. This indicates that relatively thinner substrates exhibit favorable absorption properties in the specified frequency range. A good response implies efficient interaction between EM waves and the substrate, resulting in effective absorption.

The influence of different substrate thicknesses on absorption efficiency within specific frequency ranges is illustrated in Fig. 10. When the substrate thickness is 30 μm in Fig. 10(a), absorption rates exceeding 80% are noticed between 1–2 and 3–4 THz. Conversely, in Fig. 10(b) with a thicker substrate of 40 μm , notably improved absorption performance is noticed above 90%, especially at 1.5 THz, 3 THz, and ranging from 4.5 to 5.2 THz. These findings indicate a relationship between substrate thickness and absorption capacity, with thicker substrates demonstrating absorption across the 0–6-THz frequency bands.

Fig. 11 illustrates the impact of modifying substrate thickness on absorption efficiency, with a thickness of 50 and 60 μm . In Fig. 11(a), absorption levels exceed 85% within frequency bands ranging from 0.8 to 1.5 THz, 2 to 3 THz, and at 4.5 and 5.5 THz. In contrast, Fig. 11(b) demonstrates superior absorption, exceeding 90% at 1, 1.5–2.5, 3–3.5, 4.5–5, and 5.5–6 THz. Thicker substrates enhance absorption performance, potentially by altering EM field distribution and impedance matching. These results reveal the importance of optimizing substrate thickness for specific frequency ranges essential for applications, such as sensing and communication. The study emphasizes the complex link between substrate

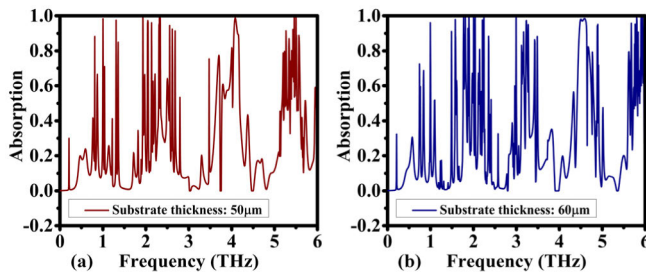


Fig. 11. Investigating the effect of the proposed substrate thickness on the ability to absorb (a) substrate thickness 50 μm and (b) substrate thickness 60 μm .

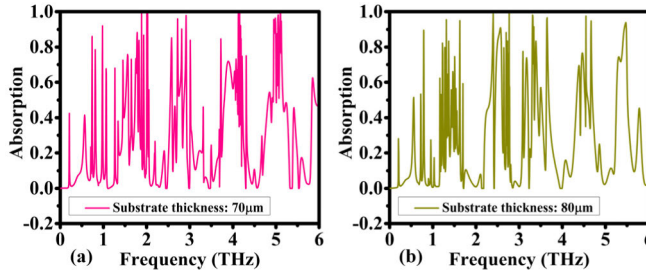


Fig. 12. Investigating the effect of the proposed substrate thickness on the ability to absorb (a) substrate thickness 70 μm and (b) substrate thickness 80 μm .

thickness and absorption capacity. Thicker substrates provide more material for EM waves to interact with, likely leading to increased absorption.

Finally, Fig. 12 further illustrates comparative study on how the thickness of a substrate impacts its ability to absorb at various frequencies in a range from 0 to 6 THz. In Fig. 12(a) where the substrate is 70 μm thick, absorption levels surpass 80% over a range of frequencies from 1 to 6 THz, showing peaks at 1, 2, 2.5–3, 4, and 5 THz, respectively. On the other hand, in Fig. 12(b) with a substrate thickness of 80 μm , absorption levels above 80% are observed within a narrower frequency span, notably at frequencies 0.5, 1–2, 3, 3.5, 4.5, and 5.5 THz, respectively. These results suggest that the thickness of the substrate significantly affects its absorption properties. Thinner substrates display wider absorption peaks spanning a broader frequency range, while thicker substrates exhibit narrower absorption peaks concentrated at specific frequencies. This understanding is crucial for optimizing substrate design for various applications, such as sensors, communication devices, and EM shielding.

D. EM Properties and Simulation Responses

Fig. 13 illustrates the impact of permeability (μ) and permittivity (ϵ) on the absorptive properties of MTMs and their interaction with EM radiation. The analysis of these components is crucial for optimizing absorption efficiency and sensitivity in detecting biological analytes. The real parts influence wave propagation and absorption, while the imaginary parts indicate the dissipative nature of the material. Furthermore, Fig. 13 demonstrates the real and imaginary

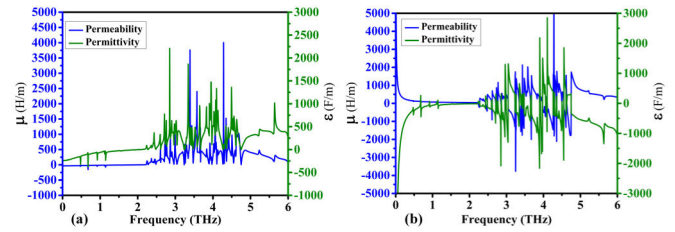


Fig. 13. (a) Real parts and (b) imaginary parts of permeability μ and permittivity ϵ of the proposed absorber.

components of the permeability (μ) and permittivity (ϵ) of the recommended absorber. Fig. 13(a) emphasizes the real parts of permeability (μ) and permittivity (ϵ), which signify the material's capacity to support H -fields and store electrical energy, respectively. These characteristics are vital for efficiently absorbing EM waves. A high real part of permeability indicates a strong magnetic response, while a significant real part of permittivity suggests the ability to store electrical energy, facilitating absorption [54]. Fig. 13(b) presents the imaginary parts of permeability (μ) and permittivity (ϵ), which demonstrate the dissipative losses in the material due to the absorption of EM energy. A high imaginary part of permeability demonstrates the effective absorption of H -field energy, while a notable imaginary part of permittivity indicates the absorption of electrical energy. These elements are crucial for ensuring the successful attenuation of incident EM waves within the material, preventing reflection or transmission [55], [56]. In addition, Fig. 13 provides insights into the interaction of the material with THz waves by examining permeability (μ) and permittivity (ϵ) across the frequency range of 0–6 THz. Through the examination of the real and imaginary parts of μ and ϵ , researchers can better understand wave propagation and absorption characteristics necessary for achieving multiband absorption tailored to specific biomolecules.

The MTM response in this article, including permeability and permittivity from 1 to 6 THz, is justified despite the unit cell periodicity of 150 μm . MTM qualities are generally specified by unit cells smaller than the wavelength; however, in the THz region, EM waves may interact with subwavelength structures to yield effective medium parameters that are suggestive of MTMs. Many resonance peaks detected within the frequency range of interest are compatible with MTM behavior caused by complicated EM interactions inside the structure. At these scales, the effective medium approximation holds true, enabling the structure to display MTM properties such as negative permeability or permittivity at certain frequencies. As a result, the periodicity limitation does not prevent the structure from being classified as an MTM within the given frequency range, allowing for an accurate interpretation of the biosensor's performance.

Fig. 14 provides a thorough evaluation of biosensor performance by analyzing impedance (z) and the reflection coefficient (S_{11}). The variations in impedance allow for the examination of the biosensor's ability to interact with biological samples and detect changes in their dielectric properties.

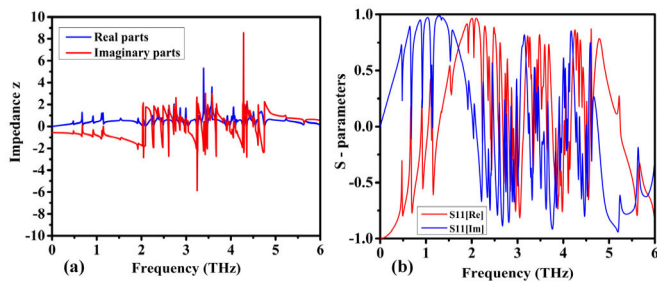


Fig. 14. Simulated responses of parameters of the proposed model (a) impedance z and (b) S_{11} real and imaginary parts.

On the other hand, the analysis of S_{11} helps to understand the biosensor's reflectance characteristics and its interaction with external stimuli. This comprehensive understanding is crucial for refining biosensor design, enhancing sensitivity, and optimizing performance for various biomedical sensing applications. Fig. 14 depicts the simulated responses of parameters in the proposed framework, with a particular focus on impedance (z) and the real and imaginary components of the S_{11} parameter. These elements play a significant role in understanding the behavior of electronic circuits, particularly in radio frequency (RF) and microwave engineering. Fig. 14(a) shows impedance (z) measures the resistance to alternating current flow in a circuit. The illustrated responses present how the circuit reacts to various frequencies. Alterations in impedance can change signal in the circuit's behaviors such as its resonance frequency or bandwidth. Understanding impedance services researcher in designing circuits that can efficiently transmit or receive signals without loss. In Fig. 14(b), the S_{11} parameter, depicted through its real and imaginary parts, is widely utilized in RF and microwave engineering to elucidate the reflection coefficient of a circuit or device. The real part denotes the reflection's magnitude, while the imaginary part signifies the phase shift resulting from the reflection. By examining both aspects of the S_{11} parameter, researcher can evaluate how effectively a circuit conveys energy from a source to a load or the amount of energy reflected due to impedance mismatches [57]. In addition, Fig. 14 evaluates absorption efficiency by examining impedance (Z) and the S_{11} parameter. A close-to-zero impedance and a magnitude of S_{11} close to 1 (0 dB) at desired absorption frequencies indicate efficient absorption of THz waves, which is essential for bio-sensing applications. These figures directly contribute to the optimization of MTM design and the prediction of sensor performance, ultimately facilitating successful biosensing within the THz range.

The proposed biosensor acts as a perfect absorber and exhibits remarkable absorption properties in the THz spectrum, specifically within the range of 0–6 THz. Its performance is distinguished by multiple sharp absorption peaks, each exceeding 95% at specific frequencies within this range. This exceptional behavior is demonstrated through the absorption spectrum, which displays several distinct peaks, as shown in Fig. 15(a). Correspondingly, the reflection spectrum shows matching dips at these same frequencies. This inverse relationship between absorption peaks and reflection dips indicates

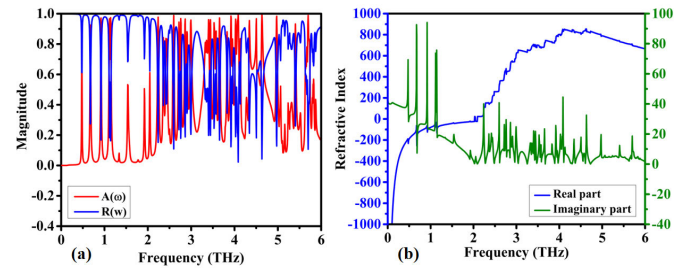


Fig. 15. Simulated responses of parameters for the proposed model (a) reflection and absorption spectra of the absorber and (b) real and imaginary parts of the refractive index.

that the absorber is highly efficient at absorbing THz radiation at these target frequencies while minimizing reflection losses. The refractive index of MTMs plays a vital role in biosensing applications, particularly for medical diagnostics, such as cervical cancer detection. To validate the proposed structure, it is essential to demonstrate that it possesses a negative refractive index and negative permittivity properties.

Fig. 13 shows the real and imaginary parts of permeability (μ) and permittivity (ϵ), which exhibit significant fluctuations and dip below zero at certain frequencies. This confirms the presence of negative refractive index properties, characteristic of MTMs, enhancing sensitivity and specificity for biosensing. Fig. 15(a) presents the reflection and absorption spectra, highlighting high absorption peaks corresponding to frequencies with negative permittivity and permeability from Fig. 13. In particular, significant absorption occurs around 2–6 THz, aligning with negative values. Fig. 15(b) shows the real and imaginary parts of the refractive index, further validating the MTM properties. The negative values at specific frequencies indicate unconventional EM wave manipulation, supporting high sensitivity and specificity. The figures demonstrate that negative permeability and permittivity (see Fig. 13) translate into practical outcomes, such as reduced reflection and increased absorption (see Fig. 15), affirming the sensor's MTM functionality. These properties ensure the sensor's capability for sensitive detection of early stage cervical cancer.

E. MTM Structure Field Distributions

Analyzing the electric field (E -field) in THz MTM perfect absorber biosensors for cervical cancer detection is vital. The real part $[\text{Re}(E)]$ indicates E -field magnitude and direction, especially the E_z -field along the z -axis, revealing energy storage and directionality. High $\text{Re}(E)$ values in MTM layers show strong THz wave interaction. The imaginary part $[\text{Im}(E)]$ reflects energy dissipation due to material properties, crucial for perfect absorption. Analyzing E_z -field with CST Studio Suite aids visualization of real and imaginary components at various frequencies, correlating with absorption characteristics (e.g., S_{11} parameter). This analysis, combined with biomarker studies, strengthens biosensor efficacy in early cancer detection at THz frequencies.

Fig. 16 displays the field distribution of an MTM perfect absorber designed for use within the 0–6-THz frequency range. The MTM absorber, characterized by a multiband response with approximately 18 resonance points, exhibits distinct

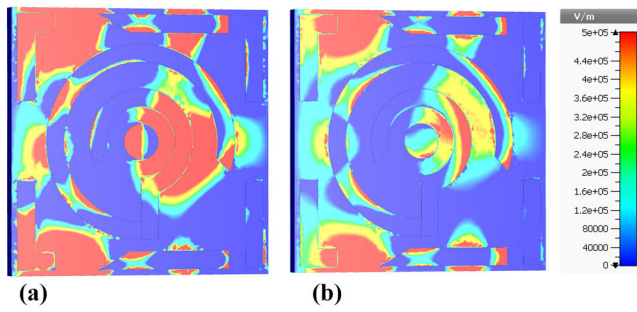


Fig. 16. Distributions of the MTM structure field are shown on a color map. (a) E_z -field real part. (b) E_z -field imaginary part.

peaks in its absorption spectrum. Fig. 16 illustrates the E -field (E_z) distributions of the MTM structure, with Fig. 16(a) representing the real part and Fig. 16(b) representing the imaginary part of the E_z -field component.

In Fig. 16(a), the real part of the E_z -field distribution is shown in a color map, where different colors represent the intensity of the E -field at various points on the MTM surface. The scale on the right, marked in V/m, indicates the E -field strength, ranging from low (blue) to high (red) values. The areas with higher field intensities correspond to the regions where the MTM structure interacts most effectively with the incident EM waves, leading to enhanced absorption.

Fig. 16(b) presents the imaginary part of the E_z -field distribution, which is critical for understanding the phase behavior of the E -field across the MTM surface. Similar to the real part, the color map indicates the field strength, with variations in the imaginary component reflecting the reactive properties of the MTM. The combination of these field distributions provides insights into the resonance behavior and efficiency of the MTM absorber, highlighting how the structure can effectively manipulate and absorb incident THz waves across multiple frequencies. This detailed analysis of the field distributions is essential for optimizing the design and enhancing the performance of the MTM absorber for applications in sensing and imaging.

Fig. 17 illustrates the E -field distributions of a THz MTM biosensor designed for early cervical cancer detection, separated into the real [see Fig. 17(a)] and imaginary [see Fig. 17(b)] components of the E -field ($/E/$ -field). The MTM structure exhibits 18 distinct sharp resonance peaks within the 0–6-THz range, demonstrating the sensor's high sensitivity and multiband capability essential for biosensing. These multiple peaks enable detailed tissue characterization, enhancing the detection of subtle changes associated with early stage cancer while also improving specificity and robustness for reliable diagnosis. The E -field density is color-mapped, ranging from 0 to 5×10^5 V/m.

In Fig. 17(a), the real part of the $/E/$ -field [$\text{Re}(E)$] demonstrates the magnitude and direction of the E -field within the MTM structure. High values, depicted in red, are predominantly observed within the aluminum resonators, signifying regions of strong interaction with THz waves and efficient absorption. This interaction is crucial for enhancing the biosensor's sensitivity to the presence of biomarkers indicative of

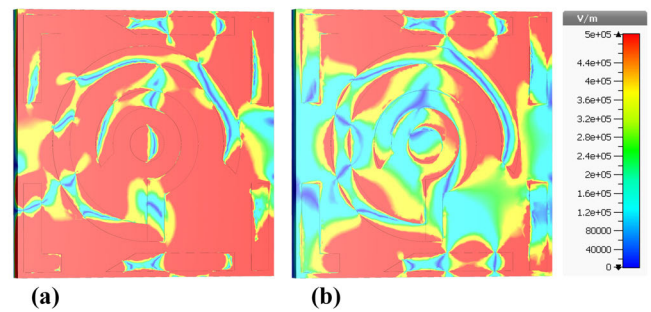


Fig. 17. Distributions of the MTM structure field are shown on a color map. (a) $/E/$ -field real part. (b) $/E/$ -field imaginary part.

cervical cancer. In contrast, Fig. 17(b) shows the imaginary part of the $/E/$ -field [$\text{Im}(E)$], highlighting the energy storage and dissipation within the MTM. Positive values, indicated by varying colors from yellow to red, suggest areas where energy absorption is efficient, aligning with the design goal of achieving perfect absorption for optimal sensor performance.

The comprehensive analysis of the $/E/$ -field components using EM simulation tools, such as CST Studio Suite allows for detailed visualization and optimization of the biosensor's performance. By identifying regions with high real part values (indicating strong field intensity) and positive imaginary part values (indicating efficient absorption), researchers can fine-tune the sensor design to maximize its effectiveness in detecting early stage cervical cancer. This approach ensures that the biosensor can accurately and reliably identify the presence of cancer biomarkers, thereby facilitating early diagnosis and improving patient outcomes.

In the context of our biosensor designed for early cancer detection, understanding the H -field's division into real and imaginary parts (H_z -field) is crucial. The real part of the H_z -field (H_z -real) is associated with the biosensor's resonant behavior and absorption properties, which directly impact its performance. Conversely, the imaginary part (H_z -imaginary) reflects energy dissipation within the biosensor due to losses, such as absorption and scattering, influencing sensitivity, and signal-to-noise ratio.

Analyzing the H_z -field for both real and imaginary parts involves utilizing software, such as CST Studio, which employs numerical methods, such as the finite element method (FEM) or finite-difference time-domain (FDTD) simulations. These simulations calculate the distribution of the H_z -field within the biosensor structure, providing insights into its behavior in the THz range (0–6 THz). CST Studio generates visualizations depicting the magnitude and direction of the H_z -field across the biosensor, with blue representing minimum intensity and red indicating maximum intensity, typically ranging from 0 to 100 A/m.

The images produced by CST Studio offer qualitative insights into how the H_z -field interacts within the biosensor at specific frequencies. Areas with stronger red hues denote regions of higher H -field intensity, providing valuable information for optimizing the biosensor's performance. However, for a comprehensive evaluation, it's essential to correlate these H_z -field characteristics, particularly the real and imaginary

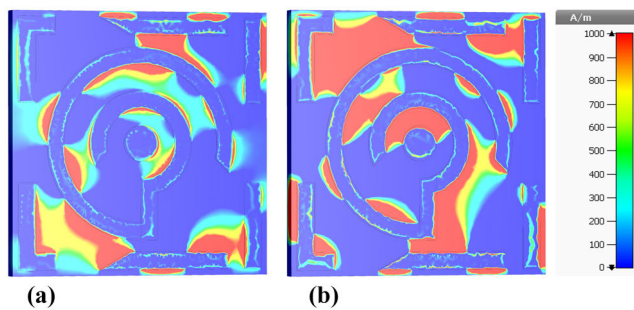


Fig. 18. Distributions of the MTM structure field are shown on a color map. (a) H_z -field real part. (b) H_z -field imaginary part.

parts, with overall absorption efficiency and resonance behavior, especially in the context of early cervical cancer detection.

Fig. 18 presents the H -field (H_z -field) distributions of an MTM-based biosensor tailored for multiband THz sensing in the 0–6-THz range, aiming at early cervical cancer detection, with H -field intensity varying from 0 to 1000 A/m. The real and imaginary parts of the H_z -field distribution are, respectively, illustrated in Fig. 18(a) and (b), playing pivotal roles in understanding the biosensor's behavior and performance. Analyzing these distributions using computational techniques allows for a detailed scientific explanation, involving calculations of field patterns, resonance frequencies, absorption spectra, and other relevant parameters, based on the MTM structure's interaction with THz waves. The color maps in Fig. 18 provide visual representations of the H -field distributions within the biosensor's MTM structure, with a color scale indicating intensity ranging from blue (0 A/m) to red (1000 A/m). Fig. 18(a) showcases a complex pattern of the H -field, with varying intensities denoted by colors, suggesting concentrated regions within the MTM structure possibly associated with resonant behavior at specific THz frequencies. In Fig. 18(b), the imaginary part of the H_z -field distribution exhibits a similar intricate pattern but with differences in intensity distribution, reflecting energy loss due to absorption or scattering and indicating the MTM's interaction with THz waves in terms of energy dissipation. The biosensor's design, as inferred from these images, is likely optimized to enhance sensitivity and specificity for early cervical cancer detection by capitalizing on the unique interactions between THz waves and the MTM structure. Its multiband nature suggests the capability to detect multiple resonant frequencies, potentially corresponding to different cancer biomarkers. Understanding both the real and imaginary parts of the H_z -field distributions is critical for refining the biosensor's design, considering factors such as MTM geometry, composition, and their interactions with THz waves, all crucial for optimal sensor performance.

Analyzing the real and imaginary part's absolute value of the H -field ($|H|$ -field) in the context of our biosensor design is crucial for understanding its performance, especially in THz frequencies. In MTM-based biosensors, the real part absolute value of the H -field $|H\text{-real}|$ relates to the distribution and strength of the H -field within the device. It characterizes the resonant behavior and absorption properties of the biosensor. A strong correlation between the real part of the H -field

and biosensor performance indicates efficient absorption and detection of cancer biomarkers.

Conversely, the imaginary part absolute value of the H -field $|H\text{-imaginary}|$ represents the dissipation of energy within the biosensor due to losses, such as absorption and scattering. Higher values of $|H\text{-imaginary}|$ signify increased energy loss, which can impact the sensitivity and signal-to-noise ratio of the biosensor. Monitoring the imaginary part of the H -field is essential for optimizing the device's performance and minimizing signal degradation.

CST Studio Suite, a robust EM simulation software, is utilized for analyzing EM field behavior within intricate structures, such as the described biosensor. In assessing the absolute value of the H -field ($|H|$ -field) for real and imaginary components, CST Studio Suite employs numerical methodologies, particularly FEM or FDTD simulations. Initially, the software generates a mesh to discretize the simulation domain, encompassing the biosensor and its environment, facilitating the numerical solution of Maxwell's equations. Subsequently, parameters such as frequency range, material properties, and excitation sources are specified in the solver setup phase. Following this, CST Studio Suite executes numerical simulations based on the defined parameters, iteratively solving Maxwell's equations to determine EM field distributions within the biosensor. Upon completion, the software produces visualizations of EM fields, including the $|H|$ -field, representing its magnitude and spatial distribution. These visualizations, outputted in formats such as 2-D plots or color maps, aid researchers in interpreting H -field behavior within the biosensor, facilitating optimization for efficient cancer biomarker detection. By visualizing $|H\text{-real}|$ and $|H\text{-imaginary}|$, researchers can assess the resonant behavior, absorption efficiency, and losses of the biosensor across different frequencies in the THz range.

Fig. 19 depicts the H -field distribution within a THz biosensor using color maps (0–2000 A/m, blue–red scale). Fig. 19(a) shows the real part ($|H\text{-real}|$), a complex pattern with swirling colors indicating the H -field intensity. Red and yellow areas represent high intensity, while blue signifies low. This part relates to energy storage and offers insights into the biosensor's resonance behavior. Fig. 19(b) shows the imaginary part ($|H\text{-imaginary}|$), also displaying a complex pattern but dominated by red, indicating higher overall intensity. This part reflects energy dissipation within the MTM due to absorption and scattering. The color distribution reveals how the biosensor interacts with THz waves. The intricate patterns suggest the MTM is designed to create localized enhancements of the H -field, potentially improving sensitivity and specificity for detecting cervical cancer biomarkers. Analyzing both parts of the H -field distribution allows researchers to refine the biosensor design for optimal performance at specific THz frequencies.

Fig. 20 illustrates the surface current distribution on an MTM structure tailored for a THz-range biosensor aimed at early cervical cancer detection. The color scale indicates current intensity from 0 A/m (blue) to 1000 A/m (red). In Fig. 20(a), the real part of the surface current distribution is depicted, showcasing arrows representing the

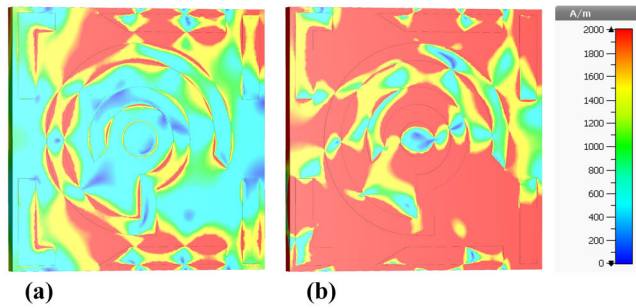


Fig. 19. Distributions of the MTM structure field are shown on a color map. (a) $H/-$ field real part. (b) $H/-$ field imaginary part.

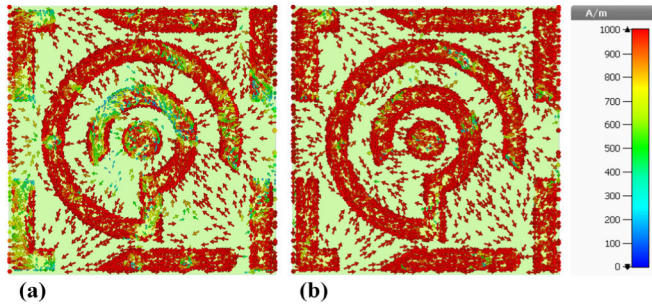


Fig. 20. Recommended MTM's surface current distribution. (a) Real part. (b) Imaginary part.

current direction and magnitude across the MTM's surface. Red arrows highlight areas of intense current, indicating strong resonant interactions with THz waves and suggesting frequency-sensitive biomarker detection. Fig. 20(b) displays the imaginary part of the surface current distribution, with red arrows denoting higher intensity. This part reflects energy dissipation within the MTM, including inherent resistance and other dissipative effects, crucial for biosensor sensitivity and specificity. The biosensor's design, evident in these images, is pivotal for performance, revealing its interaction with THz waves to detect cervical cancer biomarkers. Analyzing both parts of the surface current allows researchers to refine the biosensor's design for optimal detection capabilities across multiple THz frequencies. The ultimate objective is to develop a biosensor capable of early stage cervical cancer detection by selectively identifying molecular signatures, potentially improving patient outcomes through timely diagnosis and treatment.

IV. DIAGNOSIS OF CERVICAL CANCER

To gain more details and rigorously investigate the performance of the healthy cervical and cervical cancer sensor, here, we used numerical analysis to simulate the structure. The 3-D shape in Fig. 21 displays the MTMs' configuration of the structure. Here, the structural parameters are set as four layers including T1, T2, T3, and T2 so that the absorption spectrum is used for detection in resonance wavelength near the THz range. As can be seen, a coverslip is placed on the structure to enhance of light-matter interaction leading to higher-to-higher absorption. Besides, to obtain the same results, HeLa cell carcinoma is defined. Then, the footprint of the structure is in

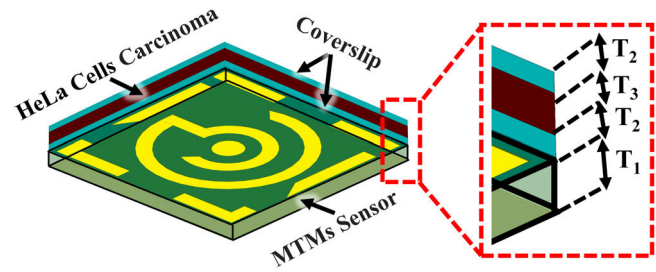


Fig. 21. Suggested biosensor investigates the absorption coefficient in both healthy cervical and cervical cancer.

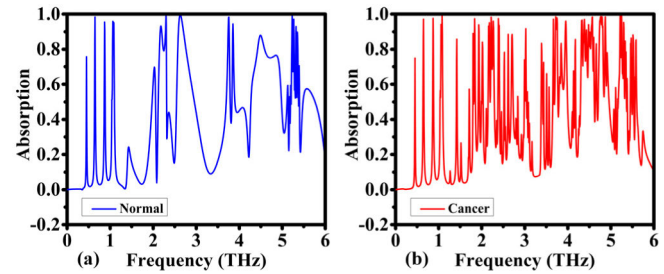


Fig. 22. Proposed biosensor detects absorption coefficients in the frequency range from 0 to 6 THz. (a) Normal cervical. (b) Cervical cancer.

agreement with the experimental point of view corresponding to the multiresonance modes.

HeLa cells, commonly used in cancer research, have a refractive index of around 1.38, with specific values of 1.392 for malignant tissue and 1.368 for healthy cervical tissue [58], [59], [60], [61], [62], [63]. Culturing HeLa cells requires a nutrient-rich medium rather than water, as water alone would cause cell death due to osmotic stress. The presence of HeLa cells influences the sensor response by altering the dielectric environment, leading to shifts in resonance peaks. This frequency dependency results in different optical properties and absorption bands. In addition, HeLa cells' unique behaviors and genetic variations over time can impact the reproducibility of experimental results, necessitating careful consideration to ensure data accuracy.

Cervical cancer is a type of cancer that occurs in the cells of the cervix, which is the lower part of the uterus that connects to the vagina. Diagnosis of cervical cancer typically involves a combination of medical history, physical examination, Pap smear, HPV testing, and biopsy. During a Pap smear, cells are collected from the cervix and examined under a microscope to look for any abnormalities. If abnormal cells are found, further testing may be done, including HPV testing to check for the presence of the human papillomavirus, which is a known risk factor for cervical cancer.

If abnormal cells are detected, a biopsy is usually performed to confirm the diagnosis of cervical cancer. A tissue sample is taken from the cervix and examined by a pathologist to determine if cancer is present.

To compare the absorption properties of the proposed sensor based on the cavity with a structure based on the circular cavity, the absorption spectrum in the normal case (blue curve) and in the detection case (red curve) is also shown in

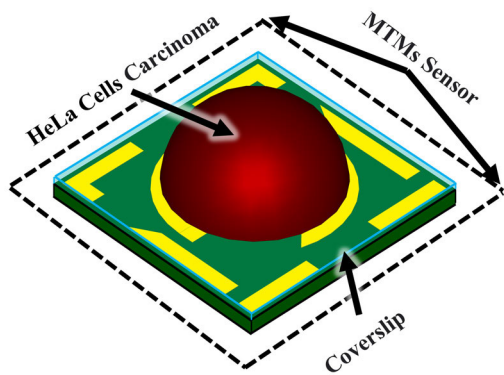


Fig. 23. Diagnosis of cervical cancer using the MWI technique.

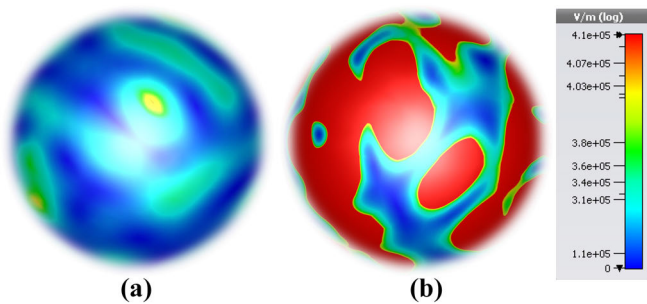


Fig. 24. E -field MWI technique results. (a) Normal cervical. (b) Cervical cancer.

Fig. 22(a) and (b), respectively. The corresponding resonance frequency occurs from 0 to 6 THz, with absorption values of approximately 1 and FWHM values of about 0.5 THz around 2–3 THz.

MWI is a developed method with noteworthy potential to supplement recognized imaging methods to enhance biomaterial and cancer detection. To date, numerical simulations have been applied to analyze the different MWI methods, but some methods have been conducted. To cope with this issue, here numerical simulation with MWI technique is a reliable scenario that can improve absorption which leads to higher sensitivity. The same structure is shown in Fig. 23.

To further investigate the performance of the above-mentioned structure, we have also inspected the E -field MWI technique in two cases: 1) the left panel (normal case [see Fig. 24(a)]) and 2) the right panel (cancer case [see Fig. 24(b)]). The E -field for these cases is tuned from 0 to 4.1×10^5 V/m. As can be seen by considering the cancer case [see Fig. 24(b)] high light–matter interaction is occurred.

H -field MWI is a technique used in nondestructive testing and imaging to detect and visualize hidden defects or anomalies in materials. It works by transmitting microwave signals and measuring the H -field response generated by interactions with the material. This technique can be used for various applications such as quality control in manufacturing, inspection of composite materials, and medical imaging.

In contrast, in order to study the relationship between the two cases with H -field considering the MWI technique, the numerical results are calculated. As can be seen in

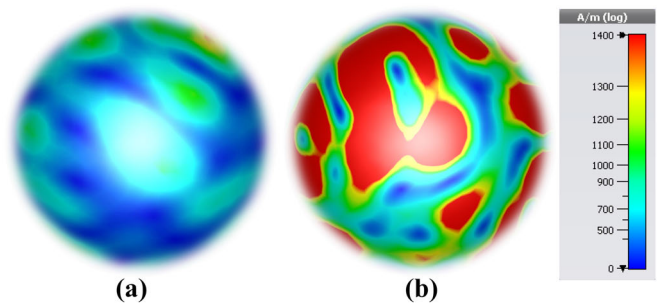


Fig. 25. H -field MWI technique results. (a) Normal cervical. (b) Cervical cancer.

TABLE II

BIOSENSING PERFORMANCE COMPARISONS OF VARIOUS SENSOR APPLICATIONS BASED ON MTM

Ref.	Year Published	FOM (RIU ⁻¹)	Q	S (THz/RIU)	Bio-application
[16]	2014	0.1216	5.58	0.02432	detection of Penicillia
[17]	2017	-	-	0.0242, 0.02438	detection of Virus
[18]	2020	-	-	0.960	Biosensor, Collagen sensor
[19]	2020	1.88	6.6	0.285	Cancer detection
[27]	2021	-	10.64	1.65	Polystyrene particle
[20]	2021	-	-	0.2833	Cervical cancer
[38]	2021	-	-	0.074	Cancer detection
[29]	2022	3.86	13	0.207	detection of avian influenza virus
[22]	2022	0.166	-	1.06	Hepatocellular carcinoma sensor
[64]	2022	-	-	0.068	Cancer Diagnosi, Biosensor
[21]	2022	1.81, 1.57	8.21, 6.05	0.203	Bovin serum albumin protein
[14]	2022	2.75	2.43	1.21	Cancer detection
[65]	2023	-	11	0.278	Non-Melanoma Skin Cancer Diagnostics
[66]	2023	-	82	0.495	
[28]	2023	0.86, 1.15	12.8, 13.5	0.0515, 0.076	
This work	-	3525838	45.056	236640	Cervical cancer

Fig. 25(a) and (b), H -field is tuned from 0 to 1400 A/m. The main result is seen in absorption and high interaction.

V. FUTURE PERSPECTIVE

Early stage detection of various cancers, including colon cancer, blood cancer, adrenal gland cancer (PC-12), breast cancer, and nonmelanoma skin cancer, can be achieved through the use of THz EM (EM) wave imaging biosensors.

VI. BENCHMARKING

This article introduces a cutting-edge biosensor engineered with an MTM perfect absorber, tailored for the early detection of cervical cancer. Operating within the THz range of 0–6 THz, the biosensor is multiband, featuring over 18 resonance peaks. The performance metrics of this biosensor are exceptional when compared to previously developed biosensors, as evidenced by its high sensitivity of 236 640 THz/RIU, high quality factor of 45.056, and an impressive figure of merit (FOM) of $3\,525\,838\text{ RIU}^{-1}$, detailed in Table II.

TABLE III
COMPARISON BETWEEN THE THz BAND STUDY ON PERFECT MTMs AND THE RECOMMENDED BIOSENSOR

Ref.	Material substrate	Frequency operating THz	Absorptivity	Techniques used	Application
[39]	SiO2	7-9.5	0.98	Au/SiO2/ Graphene	Multi-Frequency Broadband and Ultra-Broadband
[40]	silicon dioxide	1.5-1.7	0.972, 0.991	Gold/silicon dioxide/ Gold	biosensor for detecting coronaviruses
[14]	SiO2	0.5-2.5	-	SiO2/Graphene	Breast cancer detection
[41]	SiO2	2-6	0.99	graphene/Au/SiO2/Au	Refractive index sensor
[15]	Glass	0-0.37	0.998	Glass/InSb/MgF2/InSb	Colon Cancer Detection
[42]	photonic crystal plate	1-3	0.97, 0.98, 0.99	bulk Dirac semimetal/photonic crystal/Au	Narrowband perfect absorber Sensor
[43]	dielectric layer	1-3	0.99, 0.99	Au/dielectric layer/Au	
[44]	PET	0-3	0.99, 0.80, 0.95	PET/FSS/UV glue/ Graphene	Multifunctional Tunable Terahertz
[45]	Teflon	0.7-5	>0.96	Ion gel/Graphene/Teflon/Gold	polarization-sensitive
[46]	Topas spacer	0.5-4.5	0.99, 0.98, 0.99	graphene/Topas/Au	Ultra-Broadband Absorber
[47]	dielectric Teflon	1-2.2	0.99	Au/dielectric Teflon/Au	Sensor
This work	Polyimide	0-6	Ten peaks: > 0.97 Six peaks: > 0.98 Two peaks: > 0.998	Al/ Polyimide /Al	Cervical cancer, Microwave Imaging and Biosensor

Further demonstrating its superiority, the proposed biosensor achieves remarkable absorbance levels across its resonance peaks: ten peaks exceed 97% absorbance, six exceed 98%, and two surpass 99.8%, within a 6-THz operating bandwidth, as summarized in Table III. This competitive performance highlights the proposed biosensor's potential for significant advancements in early stage cervical cancer detection, outperforming many existing biosensors in both sensitivity and efficiency.

VII. CONCLUSION

This study focused on developing a novel multiband biosensor structure for the early stage detection of cervical cancer. The sensor is an MTM-based design intended to operate in the low-THz range, from 0 to 6 THz. It features an MTM layer implemented as a patterned aluminum structure placed on a polyimide substrate. The design process was meticulous, aiming to achieve near-perfect absorption of incoming EM waves across the entire operating frequency range. Full-wave EM simulation tools were used to accurately evaluate the system properties. This article provides a detailed explanation of the intermediate steps in the sensor's topology development along with an EM-based investigation of the absorption characteristics. Additional studies examined the effects of the substrate material and thickness and the resonator material. An MWI system incorporating the device was analyzed to demonstrate the proposed sensor's utility for early stage cancer diagnosis. Extensive simulation studies consistently corroborated the sensor's ability to distinguish between healthy and cancerous cervical tissue. Furthermore, benchmarking against state-of-the-art sensor designs reported in the literature highlighted the proposed sensor's remarkable performance, particularly in terms of absorbance levels and the width of the operating

bandwidth. These features make the proposed sensor a suitable tool for high-sensitivity cancer detection.

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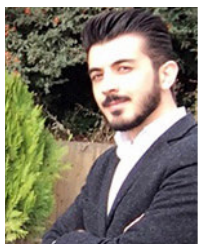
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