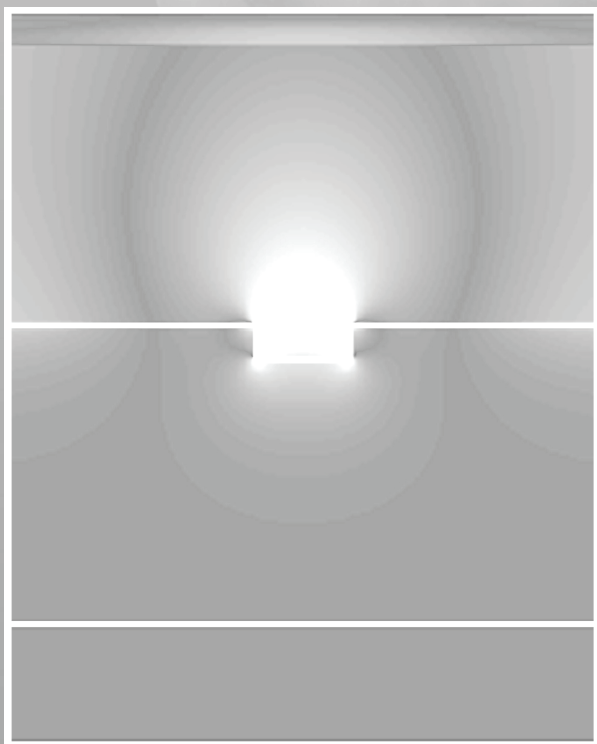


Takeo Nishikawa | Satoshi Fujita

NANOIMPRINT BIOSENSORS

**The Fusion of Nanofabrication, Nanophotonics,
and Nanobiology**





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Preface

In recent years, the correlation between the health condition of human beings and biological molecules such as nucleic acid and protein, which are major components of the human body, is becoming clearer due to advances in medical technology. Some of these biological molecules called tumor markers are being used as diagnostic indicators in medical tests. In the near future, by detecting the kind, amount, and condition of biological molecules with higher sensitivity and higher accuracy, it would be possible during preventive medical care to remedy and remove a disease before its emergence. Currently, such detection in biological molecules is difficult and costly, and implementing biosensors for the purpose will be greatly helpful in controlling health conditions, reducing medical costs significantly, and improving quality of life of human beings. However, for detecting target molecules with diameter as small as a few to a few tens of nanometers and in low concentration, biosensors with extremely high sensitivity are required. Furthermore, the sensing system should also be compact and popular priced to make its usage widespread.

We have been working to achieve these technically challenging issues by utilizing “nanotechnology” that has been keenly focused upon in this decade. In the conventional biosensing system, the detecting system has a dimension larger than millimeter scale while that of the target molecule is in nanometer scale. Therefore, many additional functions and elements are necessary to compensate the error factors derived from the difference in the detection scale in this system. This results in an increase in the apparatus size and consequently high cost. In our method, the target molecules are detected by a sensor device in which nanopatterns are formed by nanofabrication technology. The nanopatterns are made to interfere with light incident on a nanoscale on the basis of nanophotonics principle and thus ambient minute changes are detected. In this method, since the dimensions of the detection system and the target molecules are in the same order, superfluous noise factors can be

eliminated and a high signal-to-noise ratio can be realized. The size of the biosensor system can be as small as a palm in this technology. To achieve high sensitivity, we have also developed a biological probe layer on the sensor surface and it plays the most important part in the procedure which is to capture the target molecules in a specific way. In this biosensor, many different kinds of technologies are included, which have been presented in the book. We feel that fusion of the state-of-the-art technologies with nanofabrication, nanophotonics, and nanobiology fields will be the key to realize new biosensors to satisfy detection demands of the next decade.

In Chapter 1, nanofabrication technologies, especially the nanoimprint technology that can realize mass production of the devices with nanoscale patterns on it, are introduced. Although these technologies have been applied to various applications, such as recording medium, much smaller-scale patterns are necessary to use them for the detection of biological molecules. In addition, a high process throughput is also required to achieve low-cost sensor devices. Examples of basic technologies and applications are also introduced in this chapter.

In Chapter 2, behavior of light in nanoscale is described. In regions smaller than its wavelength (several hundred nanometers), light exhibits “wave” nature. As one of its interesting phenomena, it can resonate with free electrons in a metal. This phenomenon is called “surface plasmon resonance” and is applied to detect the change in ambient resonant condition. However, the resonant interaction of light with free electrons, which is confined in the nanoscale metal structure, has not been analyzed in detailed so far and is being keenly focused upon in recent years.

In Chapter 3, recent trends in biotechnology to develop a surface membrane that can be used on a sensor to capture target biomolecules is introduced. To realize a highly sensitive biosensor, the construction of a probe layer on the basis of an oligo(ethylene glycol)-mixed self-assembled monolayer is very significant. Our developments to achieve a probe layer with high sensitivity and high specificity are also included in this chapter.

In Chapter 4, we introduce the nanoimprint biosensor which we have proposed. As mentioned above, detailed analysis of resonance between light and the nanopatterned metal has not been conducted yet. In this chapter, results of the design, fabrication, and evaluation

of the sensor device using nanoimprint technology are presented along with a description of the optimization of nanopatterns for the nanoimprint biosensor.

In Chapter 5, proto-type systems for the nanoimprint biosensors are presented. We have constructed three different types of the proto-models. The first one is for laboratory use, the second for on-site use, and the third for portable use. We have developed a palm-sized biosensor system especially for portable use.

In Chapter 6, applications of nanoimprint biosensors are presented. Detection results of biological molecules using a fabricated sensor are reported. The chapter discusses in detail the use of gold nanoparticles and magnetic beads with nanoimprint biosensors to achieve higher sensitivity, although such systems are still being developed. Besides their use in medical care, application concepts of nanoimprint biosensors in other fields are also introduced in this chapter.

The technologies and concepts of nanoimprint biosensors to detect small molecules can be applied to a wide range of fields such as environmental sensing, food analysis, security check, as well as medical care and healthcare in the future. We hope that this book proves helpful for researchers working on this concept and can thus contribute to improve human life.

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Satoshi Fujita

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