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# Application of CD169, CD64 infection index in differential diagnosis of infectious diseases

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Abstract: This study focuses on the biological characteristics of CD169 and CD64 and their applications in infection diagnosis. CD169, the sialic acid receptor, plays an important role in immune response, participating in immune regulation and cell-cell interactions by recognizing and binding to pathogens that acidify. As a receptor for FC-γ, CD64 plays a key role in the immunoglobulin superfamily, mediating the phagocytosis of immune complexes and influencing immune responses. They interact in the immune response and enhance the immune effect together. Bacterial and viral detection methods, as well as the use of white blood cell count and C-reactive protein in the diagnosis of infection are also outlined, while the limitations of existing diagnostic methods are pointed out. In the application of CD169 and CD64, flow cytometry has become an important assay to quantify specific molecules on the cell surface and provide accurate information for infection diagnosis. Further study on the role of CD169 and CD64 in infection diagnosis is of great significance for improving the level of diagnosis and treatment.

#### 1. Introduction

In the field of medical research, the diagnosis and treatment of infectious diseases has been a major topic that has puzzled doctors and scientists all over the world. With the continuous variation of pathogens and the aggravation of antibiotic resistance, traditional diagnostic methods have been difficult to meet the clinical needs, and it is urgent to find new biomarkers and diagnostic strategies. In this context, the research significance of CD169 and CD64, two cell surface molecules, has become increasingly prominent. This study aims to explore the biological characteristics of CD169 and CD64 and their application value in infection diagnosis, in order to provide theoretical basis for early detection and accurate treatment of infectious diseases. Further research on the important role of CD169 and CD64 in immune response and their potential applications in infection diagnosis will not only reveal the complex mechanism of immune regulation, but also provide clinicians with more efficient and reliable diagnostic means. This is of great practical significance for improving the diagnosis and treatment level of infectious diseases, alleviating patients' pain and reducing medical costs, and also provides new ideas and directions for the research and development of new anti-infective drugs.

# 2. Biological characteristics of CD169 and CD64

#### 2.1 Biological function of CD169

CD169, also known as sialic acid receptor, is a cell surface molecule that is mainly expressed on the surface of specific immune cells, such as macrophages and dendritic cells. Its biological functions are diverse and complex, especially in immune response and cell-cell interactions play a key role<sup>[1]</sup>. CD169 plays an important role in immune surveillance and pathogen clearance. As a pattern recognition receptor, CD169 is able to specifically recognize and bind to pathogens containing sialic acid residues, such as bacteria and viruses. This recognition promotes phagocytosis and degradation of the pathogen by macrophages, thereby limiting the spread of the pathogen in the early stages of infection. In addition, this function of CD169 is also involved in the presentation of pathogen-associated molecules, laying the foundation for the activation of specific immune responses. CD169 plays an important role in antigen presentation. It is able to present phagocytic pathogens or fragments of them to T cells and activate cellular immune responses. This presentation effect is not limited to pathogens, but also includes tumor antigens and autoantigens, and therefore has a potential role in the pathogenesis of anti-tumor immunity and autoimmune diseases. CD169 also plays an important role in cell adhesion and signaling. It can participate in intercellular adhesion and affect cell migration and localization. In inflammatory response, CD169 regulates the infiltration and activation of inflammatory cells through mediating intercellular interactions, thus affecting the process of inflammatory response. CD169 also activates intracellular signaling pathways that regulate cell survival, differentiation, and function. It is worth noting that the expression of CD169 is not static, but is regulated by many factors. The expression of CD169 changes in different physiological and pathological states, and this dynamic nature makes its role in immune response more flexible and complex. For example, in states such as infection, inflammation, or tumor, the expression of CD169 is up-regulated, enhancing the activity of immune cells, while in certain immune tolerance states, its expression may be down-regulated. CD169 also plays an important role in the occurrence and development of certain diseases. In HIV infection, for example, CD169 exacerbates the spread of the virus by facilitating its capture and transmission. In some tumors, CD169 expression is associated with tumor aggressiveness and poor prognosis.

#### 2.2 Biological function of CD64

As a member of the immunoglobulin superfamily, the biological function of CD64 shows high complexity and variability in multiple levels of immune response<sup>[2]</sup>. First, CD64, as an FC-γ receptor, plays a key role in innate immunity by mediating the phagocytosis of immune complexes. In the event of pathogen invasion, CD64 can recognize and bind to the Fc segment of IgG antibodies, enhancing the phagocytosis activity of macrophages and neutrophils, which not only accelerates the clearance of pathogens, but also promotes the release of inflammatory mediators. Secondly, CD64 also plays a bridging role in adaptive immune response, influencing antibody production and immune memory formation by regulating the interaction between B cells and T cells. In addition, the expression of CD64 is affected by many factors, such as cytokines, pathogen components, etc. The plasticity of this expression makes the function of CD64 in immune response more varied. In pathological conditions, upregulation of CD64 is often associated with the severity of infectious disease and is therefore considered as a potential biomarker. To sum up, the biological function of CD64 is not only reflected in its direct immune effect, but also in its regulatory role in the immune network. The diversity and complexity of this regulatory role provide rich research connotation for in-depth understanding of the operating mechanism of the immune system.

#### 2.3 The role of both in immune response

In the complex network of immune response, the two cell surface molecules, CD169 and CD64, each play a unique and important role, and their interaction and synergistic effect are crucial for maintaining the immune balance of the body and effectively resisting pathogen invasion [3]. As a sialic acid receptor, CD169 plays an important role in immune response in the following aspects. First, as a pioneer in immune surveillance, CD169 is able to recognize and bind to sialic pathogens, promoting their phagocytosis and degradation by macrophages, thereby limiting the spread of pathogens in the early stages of infection. Second, CD169 plays a bridging role in the process of antigen presentation, presenting pathogen-related molecules to T cells, activating specific immune responses, and promoting the formation of immune memory. The third is the role of CD169 in cell adhesion and signal transduction, which can regulate the migration and activation of immune cells and affect the process of inflammatory response. As a member of the FC-y receptor of the immunoglobulin superfamily, the role of CD64 in immune response should not be ignored. The main function of CD64 is to mediate the phagocytosis of immune complexes and enhance the phagocytosis activity of macrophages and neutrophils, thus playing a key role in innate immunity. In addition, CD64 also plays an important role in adaptive immune responses, influencing antibody production and immune memory formation by regulating B cell and T cell interactions. During pathogen infection, the expression of CD64 is upregulated, which enhances the phagocytic ability of immune cells, and at the same time, amplifies the immune response by activating immune cells and releasing inflammatory mediators.

The interaction and synergistic effects of CD169 and CD64 in immune response are manifested at multiple levels. During pathogen infection, activation of CD169 can promote the up-regulation of CD64 expression, thus enhancing the phagocytosis activity of immune cells. On the contrary, the activation of CD64 can also negatively regulate the expression of CD169, forming a positive feedback loop and jointly enhancing the immune response. This interaction not only reflects the synergy of the two in immune response, but also reveals the complexity and dynamics of immune regulation mechanisms.

### 3. Overview of clinical diagnosis methods

## 3.1 Bacterial and virus detection methods

In The methods for bacterial detection mainly include traditional culture methods, biochemical reactions, immunological detection, and modern molecular biology techniques<sup>[4]</sup>. Firstly, the culture method conducts aseptic culture of patient samples, observes the characteristics of colonies, and then identifies the species of bacteria. Secondly, biochemical reactions are differentiated based on the differences in bacterial metabolites. Once again, immunological detection utilizes the combination of specific antibodies with bacterial antigens and realizes the rapid detection of bacteria through methods such as enzyme-linked immunosorbent assay (ELISA). And modern molecular biology techniques, such as polymerase chain reaction (PCR) and its derivative techniques, can accurately identify bacteria at the genetic level. Compared to bacterial detection, viral detection methods are more complex and diverse. On the one hand, virus particles are tiny and difficult to observe through conventional optical microscopes. On the other hand, the viral genome has high variability, presenting challenges for detection. Currently, the main methods for viral detection include electron microscopy observation, virus isolation and culture, serological detection, and molecular biology techniques. Electron microscopy observation can directly observe the morphology of viruses, but the operation is complex and costly. Although virus isolation and culture can obtain virus strains, it is time-consuming and inefficient. Serological detection, such as neutralization tests and immunofluorescence assays, determines the situation of viral infection by detecting virus-specific antibodies in the patient's serum. And molecular biology techniques, especially real-time fluorescence quantitative PCR (qPCR) and high-throughput sequencing technology, have become research hotspots in the field of viral detection. They can not only achieve rapid and accurate detection of viruses but also monitor virus variations, providing an important basis for epidemic prevention and control and vaccine research and development. In conclusion, the continuous development of bacterial and viral detection methods provides a wealth of means for clinical diagnosis and helps improve the level of disease diagnosis and treatment.

## 3.2 The application of white blood cell count and C-reactive protein (CRP)

In the field of diagnosing infectious diseases, the application of white blood cell count and C-reactive protein (CRP) is like two mirrors, each reflecting the subtle changes in the immune status of the body. White blood cell count, as a traditional and widely used indicator, provides initial clues for determining the presence or absence of inflammation by measuring the total number of white blood cells in the blood. The increase or decrease of different white blood cells such as neutrophils, lymphocytes, and monocytes each reveals the specific tendencies of the infection type and immune response. Meanwhile, CRP, as an acute-phase protein, its elevation in the state of infection or inflammation is like a signal flag, indicating that the body is actively responding to external invasion. The application of CRP lies not only in its ability to respond quickly to inflammatory changes but also in its ability to reflect the degree and dynamic process of inflammation. The application of white blood cell count and CRP is not isolated. Their value in clinical diagnosis often complements and validates each other. In some cases, the white blood cell count may be normal while CRP has already increased, or vice versa. This variability and inconsistency require clinicians to have a high degree of complexity and flexibility when interpreting laboratory results. In addition, the influence of factors such as age, genetic factors, and chronic diseases also adds uncertainty to the application of white blood cell count and CRP. Therefore, how to accurately grasp the application of these two indicators in the complex clinical situation is not only a challenge for clinicians but also the key to improving the accuracy of diagnosing infectious diseases.

### 3.3 Limitations of existing methods

In the diagnosis practice of infectious diseases, although the existing detection methods such as white blood cell count and C-reactive protein determination have been widely used in clinical practice, their limitations should not be ignored. First, the sensitivity and specificity of these methods are not infallible, and they may have false positive or false negative results in some cases, and this uncertainty affects the accuracy of the diagnosis to some extent. Secondly, traditional indicators such as white blood cell count lack accurate differentiation of infection types and can not reveal the specific pathogen of infection, so it is powerless to guide treatment. Moreover, the varying degrees of standardization of existing methods, differences between laboratories, and differences in testing methods can cause fluctuations in results, which adds additional complexity to clinical decision making. In addition, for some special populations, such as immunosuppressed patients or the elderly, changes in these routine indicators may not be significant, leading to delayed diagnosis or misdiagnosis. At the same time, the development of infectious diseases is a dynamic process, and the existing detection methods often can only provide static information at a certain point in time, and it is difficult to fully reflect the evolution of the disease. More importantly, these approaches are particularly inadequate in dealing with emerging pathogens and antibiotic resistance. Therefore, it must be recognized that the limitations of existing diagnostic methods are the bottleneck in the diagnosis and treatment of infectious diseases, which urgently requires the medical community to explore more accurate and efficient diagnostic strategies in order to provide better medical services for patients.

# 4. Application of CD169 and CD64 in infection diagnosis

#### 4.1 The role of CD169 in the diagnosis of viral infection

In the field of diagnosis of viral infection, CD169 functions like a sharp scalpel, accurately cutting into the kernel of pathogen identification. As a cell surface receptor, CD169 reveals subtle clues to viral infection through its specific binding to viral surface sialic acid molecules. In the early stages of viral infection, the up-regulated expression of CD169 not only accelerates the endocytosis and degradation of virions, but more importantly, it provides a sensitive biological marker for clinical diagnosis. The application of CD169 in the diagnosis of viral infection is mainly reflected in its ability to identify specific virus strains, which is particularly significant in the diagnosis of pathogens such as influenza virus and HIV.

The function of CD169 is not isolated, and its performance in the diagnosis of viral infection shows complex variability. On the one hand, the expression of CD169 is affected by many factors such as virus type, infection stage, and host immune status, which makes its application in diagnosis more challenging. On the other hand, the dynamic change of CD169 in viral infection requires clinicians to have a high degree of insight and clinical experience when interpreting test results. Despite these complexities, the application of CD169 in the diagnosis of viral infection is still of irreplaceable value, which provides the possibility for early detection and rapid diagnosis of viral diseases, and also lays the foundation for further research on the mechanism of virus-host interaction. Therefore, it is of great significance to explore the role of CD169 in the diagnosis of viral infection.

# 4.2 The application of CD64 in the diagnosis of bacterial infection

The field of bacterial infection diagnosis, CD64 as a key immune receptor, its application value is like a beacon signal, illuminating the dark night of pathogen recognition. CD64, a member of the FC-γ receptor family, greatly enhances the phagocytic ability of phagocytic cells to bacteria through its binding to the IgGFc segment of the bacterial antigen-antibody complex, thus playing a crucial role in the early diagnosis of bacterial infections. During bacterial infection, the expression of CD64 is up-regulated, which not only reflects the body's emergency response to pathogens, but also becomes an important indicator to judge the severity and prognosis of bacterial infection.

The application of CD64 in the diagnosis of bacterial infections has not been smooth sailing. Fluctuating expression levels, individual differences, and cross-pollination with other inflammatory responses pose challenges for diagnostic accuracy. In addition, the dynamic change of CD64 in bacterial infection requires clinicians to consider the comprehensive influence of multiple factors such as infection type, pathogen characteristics and host immune status when interpreting its expression pattern. Nevertheless, the application of CD64 in the diagnosis of bacterial infections still has significant advantages, it can provide clinicians with rapid, direct infection information, which can help in the timely development of treatment strategies. Therefore, it is of great scientific and practical significance to study the mechanism of CD64 in the diagnosis of bacterial infection, optimize its detection methods and clinical applications, and improve the diagnosis and treatment of bacterial diseases.

# 4.3 Application of flow cytometry in detection

In the technical spectrum of infection diagnosis, flow cytometry has become an important means of CD169 and CD64 detection with its unique advantages. Through its high-precision cell analysis capabilities, flow cytometry is able to quickly and accurately quantify specific molecules on the cell surface, providing an accurate measure of the expression levels of CD169 and CD64. On this technology platform, the detection of CD169 and CD64 not only realizes the highly sensitive identification of infection status, but also reveals the dynamic changes of immune cells during infection through detailed analysis of cell populations.

The application of flow cytometry not only shows its complexity in technology, but also reflects changes in data interpretation. Through multi-parameter detection of cells, it comprehensively considers cell size, shape, surface markers and other characteristics, making the detection of CD169 and CD64 no longer single, but integrated into a multi-dimensional analytical framework. In this process, the application of flow cytometry requires not only the high technical level of the operator, but also the ability of the investigator to have a deep understanding of the immunological principles and data analysis. Despite the complexity of the procedure and the challenges in interpreting the results, the application of flow cytometry in the diagnosis of infection remains irreplaceable. It provides clinicians with fast and reliable diagnostic information, and provides strong support for patients' timely treatment and disease monitoring. Therefore, the application of flow cytometry in the detection of CD169 and CD64 not only promotes the development of infection diagnosis technology, but also provides a new perspective and tool for immunological research.

#### 5. General Junction

This study systematically describes the biological characteristics of CD169 and CD64, their roles in immune response and their application value in infection diagnosis. By dissecting the functions of these two cell surface molecules, we recognize their critical role in immune regulation and pathogen recognition. At the same time, this study Outlines the bacterial and viral detection methods, as well as the application of white blood cell count and C-reactive protein in clinical diagnosis, pointing out the limitations of existing diagnostic methods. Further, we discuss the important role of flow cytometry in the detection of CD169 and CD64, which provides a new idea for the early diagnosis of infectious diseases. Overall, the research results of this study provide a theoretical basis for the diagnosis and treatment of infectious diseases, and help to improve the accuracy of clinical diagnosis and treatment effect. In future studies, we need to further optimize the detection method and explore the mechanism of action of CD169 and CD64 in more infectious diseases, in order to provide more strong support for clinical practice and the development of new anti-infective drugs.

#### References

- [1] Zhang Y X, Yang Q T, Zhang M X, et al. Expression and clinical value of nCD64 and mCD169 in patients with early acute infection [J]. International Journal of Laboratory Medicine, 2023, 44(20):2453-2456.
- [2] Ikezumi Y, Kondo T, Matsumoto Y, et al. Analysis of CD169 (Sialoadhesin)-Positive Activated Macrophages in Kidney Damage in Lupus Nephritis[J]. Journal of the American Society of Nephrology, 2023, 34(11S):596.
- [3] Klaas M, Dubock S, Ferguson D J P, et al. Sialoadhesin (CD169/Siglec-1) is an extended molecule that escapes inhibitorycis-interactions and synergizes with other macrophage receptors to promote phagocytosis[J]. Glycoconjugate Journal, 2023, 40(2):213-223.
- [4] Zhang Xiaolin, Mao Qinghua, Ju Guanjun, et al. Relationship between CD64 index, sCD14, sCD163 and severe pulmonary infection after thoracoscopic radical resection of lung cancer [J]. Progress in Modern Biomedicine, 2024(002): 024.