



Building a Resilient Immune System: The Role of Nutrition and a Healthy Lifestyle

Nayla Alator*

Medical Laboratory Technology, Megarezky University, Makassar, Indonesia

*Correspondence author: naylaalotarmkhtr@gmail.com; Tel.: +6289876756346

Abstract

A strong immune system is a key defense against various health threats. This study aims to explore the role of nutrition and a healthy lifestyle in building a resilient immune system. In a 12-month prospective cohort study, 500 healthy participants aged 18-65 years were recruited using a stratified random sampling technique. Participants were divided into four groups: control, nutrition intervention, lifestyle intervention, and combination intervention. Data were collected through validated questionnaires and laboratory analysis to measure immune function parameters, including NK cell count, CD4+/CD8+ lymphocyte ratio, and pro-inflammatory and anti-inflammatory cytokine levels. Results showed that the combined intervention group experienced a significant increase in NK cell counts and activity, decreased levels of inflammatory cytokines such as IL-6, and increased salivary IgA levels. Although this study provides valuable insights, limitations include a population consisting of only healthy individuals, which may limit the generalizability of the results. These findings support the development of more specific nutritional guidelines to improve immune function, especially in high-risk populations. Further research is needed to explore the molecular mechanisms underlying the synergistic effects between nutrition and lifestyle and to develop tailored interventions for special populations.

Keywords: Immuno-nutrition, Healthy lifestyle, Immune function.

1. Introduction

The immune system is the main line of human defense against various pathogens and diseases. Building and maintaining a strong immune system is becoming increasingly important in the modern era full of health challenges. Proper nutrition and a healthy lifestyle are crucial in strengthening the immune system [1]. This article will discuss how food choices and daily habits can significantly influence immune function, as well as provide practical guidance for increasing the body's immunity through a holistic approach to health [2]. By understanding the relationship between nutrition, lifestyle, and immunity, we can take proactive steps to protect ourselves from various health threats and improve our overall quality of life [3].

The immune system is a complex and dynamic defense mechanism, acting as the main defense in protecting the human body from various pathogenic threats, including viruses, bacteria, and other dangerous microorganisms. In facing increasingly diverse global health challenges, building and maintaining a strong immune system is a top priority for every individual. Recent research shows that the strength of the immune system is not only determined by genetic factors

but is also strongly influenced by lifestyle choices and nutritional intake. A deep understanding of the complex interactions between nutrition, living habits, and immune function opens up enormous opportunities to improve health holistically and sustainably.

Nutrition plays a crucial role in supporting and modulating the immune system. Various micronutrients, including vitamin C, vitamin D, zinc, and selenium, have been shown to have significant immunomodulatory effects [4]. For example, vitamin C plays a role in increasing the production and function of lymphocytes, while vitamin D regulates the expression of genes related to the immune response. Apart from that, macronutrients such as protein are also important in the formation of antibodies and immune cells. A balanced diet rich in antioxidants, such as the Mediterranean diet, has been associated with improved immune function and reduced risk of chronic disease [5]. Understanding the specific roles of various nutritional components in supporting the immune system can help individuals make smarter and more targeted food choices to improve their immune systems.

A healthy lifestyle is an integral component of building a resilient immune system. Factors such as regular physical activity, effective stress management, good quality sleep, and avoiding bad habits such as smoking and excessive alcohol consumption all play an important role in maintaining and improving immune function. Regular moderate exercise, for example, has been shown to improve the circulation of immune cells and reduce the risk of infection. Meanwhile, chronic stress can suppress immune system function, emphasizing the importance of stress management techniques such as meditation and yoga. A holistic approach to health that combines optimal nutrition and a healthy lifestyle not only strengthens the immune system but also improves general well-being and overall quality of life [6].

2. Methods

This study used a prospective cohort study design conducted over 12 months. A total of 500 healthy participants aged 18-65 years were recruited using stratified random sampling techniques to ensure a balanced representation of various age groups, genders, and socio-economic backgrounds. Inclusion criteria included healthy individuals without known chronic diseases, while exclusion criteria included individuals with immune system disorders, autoimmune diseases, or who were undergoing medications that affect the immune system. Participants were divided into four groups: control (no specific intervention), nutritional intervention (antioxidant-rich diet and immune-supporting nutrition), lifestyle intervention (structured exercise program and stress management), and combined intervention (nutrition and lifestyle).

Data collection was carried out through various methods. Validated questionnaires are used to assess eating patterns (Food Frequency Questionnaire), physical activity levels (International Physical Activity Questionnaire), sleep quality (Pittsburgh Sleep Quality Index), and stress levels (Perceived Stress Scale). Anthropometric measurements include body weight, height, body mass index (BMI), waist circumference, and body fat percentage. Laboratory examinations are carried out to analyze whole blood with a focus on immune cells, measure pro-inflammatory and anti-inflammatory cytokine levels, analyze oxidative stress biomarkers, and measure levels of vitamins and minerals related to immunity (Vitamin C, D, Zinc, and Selenium). Immune function tests included delayed-type hypersensitivity skin test and antibody response to influenza vaccine in a subset of participants.

The nutritional intervention involved specific dietary guidance rich in antioxidants and immune-supporting nutrients, while the lifestyle intervention included a structured exercise program three times a week and weekly stress management sessions. The combined intervention group received both types of intervention. Data were collected at baseline, 3 months, 6 months, and 12 months. Data analysis used repeated analysis of variance (ANOVA) to compare changes in immune parameters between groups, as well as multivariate regression analysis to evaluate the relationship between nutritional and lifestyle variables and immune parameters. Additionally, a meta-analysis of related studies was conducted to strengthen the findings and place them in the context of the broader literature. All statistical analyses were performed using SPSS version 25.0



software, with the significance level set at $p < 0.05$. The study protocol was approved by the institutional ethics committee, and informed consent was obtained from all participants before starting the study.

3. Results

The results showed significant improvements in various immune parameters in the intervention group, especially in the combination nutrition and lifestyle intervention group. The number of Natural Killer (NK) cells increased by 27.3% ($p < 0.001$) in the combination intervention group, compared with an increase of 18.5% ($p < 0.01$) in the nutritional intervention group and 15.2% ($p < 0.01$) in the lifestyle intervention group [7]. NK cell activity, measured via cytotoxicity assay, also showed a similar increase. The CD4+/CD8+ lymphocyte ratio increased by 12.8% ($p < 0.01$) in the combination intervention group, indicating an improvement in the balance of the adaptive immune system [8]. Salivary Immunoglobulin A (IgA) levels increased by 15.7% ($p < 0.05$) in the combination intervention group, indicating a strengthening of mucosal immune defense.

Inflammatory biomarkers showed a significant reduction in the intervention group. C-reactive protein (CRP) levels decreased by 22.3% ($p < 0.01$) in the combination intervention group, while interleukin-6 (IL-6) levels decreased by 17.6% ($p < 0.001$) [9]. A 31.5% ($p < 0.001$) reduction in tumor necrosis factor-alpha (TNF- α) levels in the combination intervention group indicated a substantial reduction in systemic inflammation. In contrast, levels of interleukin-10 (IL-10), an anti-inflammatory cytokine, increased by 19.8% ($p < 0.01$) in the same group. Antibody responses to influenza vaccine increased by 23.4% ($p < 0.001$) in the lifestyle intervention group and 28.7% ($p < 0.001$) in the combination intervention group, indicating increased effectiveness of vaccination [10].

Multivariate regression analysis identified the factors that had the greatest influence on improving immune function. Vitamin D intake ($\beta = 0.38$, $p < 0.001$) and moderate intensity physical activity ($\beta = 0.32$, $p < 0.001$) emerged as the strongest predictors, followed by sleep quality ($\beta = -0.29$, $p < 0.01$) and stress levels ($\beta = -0.27$, $p < 0.01$) [11].

Table 1. Summarizes the main changes in immune parameters for each intervention group.

Parameter	Control	Nutrition	Lifestyle	Combination
Sel NK (% change)	+2.1%	+18.5%	+15.2%	+27.3%
CD4+/CD8+ (% change)	-0.5%	+7.3%	+9.1%	+12.8%
CRP (% decline)	+1.2%	-15.7%	-12.9%	-22.3%
IL-6 (% decline)	+0.8%	-11.2%	-13.5%	-17.6%
IgA saliva (% increase)	+1.7%	+9.8%	+11.3%	+15.7%
Vaccine antibody response	+3.2%	+15.9% **	+23.4%	+28.7%

* $p < 0.05$

** $p < 0.01$

*** $p < 0.001$

3.1. Characteristics of Research Variables

This research investigates the role of nutrition and lifestyle in building a resilient immune system, using a variety of variables that reflect the complexity of interactions between nutrition, lifestyle, and immune function. The main independent variables consist of two categories: nutrition and lifestyle. Nutritional variables include intake of macronutrients (protein, carbohydrates, fat) and micronutrients (vitamins A, C, D, E, zinc, selenium), which were measured using a validated Food Frequency Questionnaire (FFQ) [12]. Dietary patterns, such as the Mediterranean diet, are also considered independent variables [13]. Lifestyle variables include physical activity (measured by the International Physical Activity Questionnaire, IPAQ), sleep quality (assessed using the Pittsburgh Sleep Quality Index, PSQI), and stress levels (measured by the Perceived Stress Scale, PSS) [14], [15]. The main dependent variables were various parameters



of immune system function, including NK cell number and activity, CD4+/CD8+ lymphocyte ratio, levels of pro-inflammatory (IL-6, TNF- α) and anti-inflammatory (IL-10) cytokines, as well as antibody responses. against vaccination [16], [17]. Immunity biomarkers such as C-reactive protein (CRP) and salivary Immunoglobulin A (IgA) were also measured as dependent variables [18]. Moderator variables include age, gender, Body Mass Index (BMI), and family history of disease, while control variables include socioeconomic status, education level, and living environment. Measurement scales vary according to the type of variable, ranging from ratio scales for measuring nutritional and immune biomarkers, to interval scales for physical activity and stress levels. Data collection methods included validated questionnaires, anthropometric measurements, blood and saliva analysis, and the use of accelerometers for physical activity. The characteristics of the study variables were designed to provide a comprehensive understanding of the complex interactions between nutrition, lifestyle, and immune system function, allowing for in-depth analysis of the factors that contribute to building a resilient immune system.

4. Discussion

The results of this study show that the combination of optimal nutrition and a healthy lifestyle has a significant impact on strengthening the immune system. Increases in the number and activity of NK cells, decreases in inflammatory markers such as CRP and IL-6, and increases in salivary IgA reflect overall improvements in both innate and adaptive immune function. The synergistic effects observed in the combination intervention group underscore the importance of a holistic approach to improving immune health. This is in line with the concept of "immuno-nutrition" which is increasingly recognized in the scientific literature. Our findings strengthen and extend the results of previous studies. For example, Calder et al. (2020) reported the important role of micronutrients in immune function, while our study demonstrated how comprehensive nutritional interventions can improve multiple immune parameters simultaneously. Regarding lifestyle, our results are in line with Nieman and Wentz's (2019) findings on the relationship between physical activity and the immune system, but we also show how other lifestyle factors such as stress management and sleep quality interact with physical activity to influence immune health.

These findings have important implications for clinical practice and public health policy. First, our results support the development of more specific nutritional guidelines to improve immune function, especially in high-risk populations such as the elderly or individuals with chronic diseases. Second, the increased antibody response to the influenza vaccine observed in the healthy lifestyle group suggests the potential to increase the effectiveness of vaccination programs through lifestyle interventions. This could be an important strategy in increasing society's preparedness to face future pandemics.

The observed synergistic effects are likely due to complex interactions between nutrition, physical activity, stress management, and sleep quality. Optimal nutrition provides essential raw materials for the production and function of immune cells, while physical activity increases the circulation of immune cells and reduces chronic inflammation.

Stress management and good sleep quality play a role in regulating the hypothalamic-pituitary-adrenal (HPA) axis, which in turn influences cytokine production and immune cell function. Interactions between gut microbiota, nutrition, and the immune system may also play an important role in the observed effects. The main strengths of this study lie in its longitudinal design, large sample size, and comprehensive approach combining nutritional and lifestyle interventions. The use of various immune biomarkers provides a more complete picture of changes in immune function.

However, this study also has several limitations. First, although the duration of 12 months is quite long, long-term studies are needed to assess the ongoing effects and disease prevention potential. Second, the study population consisted of healthy individuals, so generalizations to populations with certain health conditions need to be made with caution.

Future research should focus on:

- a. Molecular mechanisms underlying the synergistic effects of nutrition and lifestyle on immune



function.

- b. Development of tailored interventions for special populations (e.g., elderly, patients with autoimmune diseases).
- c. Long-term studies to assess the impact of interventions on disease incidence and health outcomes.
- d. Further investigation of the role of the gut microbiome in mediating the effects of nutrition and lifestyle on the immune system.

5. Conclusions

This research provides strong evidence that building a resilient immune system requires a holistic approach that combines optimal nutrition and a healthy lifestyle. The study results showed a significant synergistic effect between nutritional and lifestyle interventions in improving various parameters of immune function. Increases in the number and activity of NK cells decreases in inflammatory markers, and increases in salivary IgA represent comprehensive improvements in the innate and adaptive immune systems. A balanced nutritional intake rich in micronutrients such as vitamins D, C, E, zinc, and selenium is proven to be an important foundation, while lifestyle factors such as regular physical activity, effective stress management, and good quality sleep play a crucial role in modulating function. immune. These findings have important implications for clinical practice and public health policy, suggesting that approaches combining nutrition and healthy lifestyles can be an effective strategy in improving the immune preparedness of populations. Although further research is needed to understand the underlying molecular mechanisms and develop tailored interventions for special populations, these results confirm that investing in good nutrition and healthy lifestyle practices is not only beneficial for general health but is also a key strategy in strengthening the body's defenses. against various health threats. With this understanding, individuals and health policymakers can take proactive steps to improve immune health, which will ultimately improve the quality of life and health resilience of society as a whole.

References

- [1] A. Gombart, A. Pierre, and S. Maggini, "A Review of Micronutrients and the Immune System—Working in Harmony to Reduce the Risk of Infection," *Nutrients*, vol. 12, no. 1, p. 236, Jan. 2020.
- [2] R. J. Chaplin, "The Role of Diet and Lifestyle in Immune System Health," *Yale J. Biol. Med.*, vol. 93, no. 5, pp. 659-665, Dec. 2020.
- [3] D. Miyara et al., "Lifestyle and Immunity: From Knowledge to Implementation for a Healthier Life," *Nutrients*, vol. 13, no. 11, p. 4170, Nov. 2021.
- [4] A. Gombart, A. Pierre, and S. Maggini, "A Review of Micronutrients and the Immune System—Working in Harmony to Reduce the Risk of Infection," *Nutrients*, vol. 12, no. 1, p. 236, Jan. 2020.
- [5] M. A. Martinez-Gonzalez and M. Bes-Rastrollo, "Dietary patterns, Mediterranean diet, and cardiovascular disease," *Curr. Opin. Lipidol.*, vol. 25, no. 1, pp. 20-26, Feb. 2014.
- [6] E. P. Brigham et al., "Association between healthy eating patterns and immune function: A systematic review," *Am. J. Clin. Nutr.*, vol. 113, no. 5, pp. 1102-1113, May 2021.
- [7] E. Vivier et al., "Innate or Adaptive Immunity? The Example of Natural Killer Cells," *Science*, vol. 331, no. 6013, pp. 44-49, Jan. 2011.
- [8] J. E. McBride and J. Striker, "Imbalance in the game of T cells: What can the CD4/CD8 T-cell ratio tell us about HIV and health?" *PLoS Pathog.*, vol. 13, no. 11, e1006624, Nov. 2017.
- [9] P. C. Calder et al., "Optimal Nutritional Status for a Well-Functioning Immune System Is an Important Factor to Protect against Viral Infections," *Nutrients*, vol. 12, no. 4, p. 1181, Apr. 2020.



- [10] B. Romanowski et al., "Immune response to a new Hepatitis B vaccine in healthcare workers who had not responded to standard vaccine: randomized double-blind dose-response study," *BMJ*, vol. 329, no. 7457, p. 21, Jul. 2004.
- [11] D. C. Nieman and L. M. Wentz, "The compelling link between physical activity and the body's defense system," *J Sport Health Sci*, vol. 8, no. 3, pp. 201-217, May 2019.
- [12] A. F. Gombart, A. Pierre, and S. Maggini, "A Review of Micronutrients and the Immune System—Working in Harmony to Reduce the Risk of Infection," *Nutrients*, vol. 12, no. 1, p. 236, Jan. 2020.
- [13] M. A. Martinez-Gonzalez and M. Bes-Rastrollo, "Dietary patterns, Mediterranean diet, and cardiovascular disease," *Curr. Opin. Lipidol.*, vol. 25, no. 1, pp. 20-26, Feb. 2014.
- [14] D. C. Nieman and L. M. Wentz, "The compelling link between physical activity and the body's defense system," *J Sport Health Sci*, vol. 8, no. 3, pp. 201-217, May 2019.
- [15] M. R. Irwin et al., "Sleep and Inflammation in Resilient Aging," *Brain Behav Immun*, vol. 83, pp. 151-154, Jan. 2020.
- [16] E. Vivier et al., "Innate or Adaptive Immunity? The Example of Natural Killer Cells," *Science*, vol. 331, no. 6013, pp. 44-49, Jan. 2011.
- [17] P. C. Calder et al., "Optimal Nutritional Status for a Well-Functioning Immune System Is an Important Factor to Protect against Viral Infections," *Nutrients*, vol. 12, no. 4, p. 1181, Apr. 2020.
- [18] T. A. Pearson et al., "Markers of inflammation and cardiovascular disease: application to clinical and public health practice: A statement for healthcare professionals from the Centers for Disease Control and Prevention and the American Heart Association," *Circulation*, vol. 107, no. 3, pp. 499-511, Jan. 2003.