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INTEGRATING NATURAL PRODUCT-BASED THERAPEUTICS WITH AI-DRIVEN LUNG CANCER PREDICTION MODELS

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Abstract: Lung cancer is one of the leading causes of cancer-related deaths globally, with early detection and accurate prediction playing a critical role in improving patient outcomes [1]. The healthcare industry has increasingly turned to machine learning (ML) and artificial intelligence (AI) to help identify at-risk individuals by analyzing vast amounts of data, including clinical, lifestyle, and environmental factors [2]. While AI models have shown promise in predicting lung cancer risk, their effectiveness can be limited by the breadth of data they consider [3]. To enhance these models and improve their predictive accuracy, integrating natural product-based therapeutics offers a promising solution [4]. Natural products, derived from plants, fungi, and microorganisms, have long been recognized for their medicinal properties [5]. Many of these compounds have demonstrated anti-cancer effects, including anti-inflammatory, antioxidant, and apoptosis-inducing properties [6]. Despite their known benefits, the role of natural products in lung cancer prevention and treatment remains underexplored in AI-driven predictive models [7]. This white paper presents a framework to integrate natural product data into AI-based lung cancer prediction models, offering an innovative, interdisciplinary approach to cancer risk management and prevention [8]. By incorporating natural product data into machine learning algorithms, we can create a more comprehensive and personalized approach to predicting lung cancer risk [9]. This integration will allow healthcare providers to not only assess genetic, environmental, and lifestyle factors but also consider the therapeutic potential of natural products in reducing cancer risk [10]. Moreover, these enhanced AI models could offer personalized prevention strategies, such as recommending specific natural products or lifestyle modifications based on an individual's unique risk profile [11]. This white paper outlines the benefits of this integration, including improved predictive accuracy, the ability to offer personalized recommendations, and the potential to intervene early in atrisk populations [12]. We also explore how incorporating natural product data could lead to more cost-effective healthcare by preventing or slowing the progression of lung cancer through early intervention [13]. Ultimately, this integrated approach could help transform the way healthcare providers predict and manage lung cancer risk, leading to better patient outcomes and a reduction in the global cancer burden [14].

The synergy between AI-driven predictive models and natural product-based therapeutics presents an exciting opportunity to innovate lung cancer prevention strategies, paving the way for more individualized, precise, and effective approaches in managing cancer risk [15]. By leveraging AI to enhance our understanding of how natural products interact with cancer biology, we can not only improve prediction accuracy but also offer proactive, personalized care that could save lives and reduce long-term healthcare costs [16].

Keywords: Artificial Intelligence, Machine Learning, Cancer, Prediction, Natural Compounds, Therapeutics, Environment

1. Introduction

Lung cancer is one of the most pervasive and deadly cancers worldwide, responsible for a significant number of cancer-related deaths each year. Early detection plays a crucial role in improving survival rates, but despite advances in diagnostic technologies, lung cancer often goes undiagnosed until it reaches an advanced stage, making treatment options more limited. Early intervention and accurate risk prediction are, therefore, critical to reducing mortality and improving patient outcomes. Over the past decade, machine learning (ML) and artificial intelligence (AI) have emerged as transformative technologies in healthcare, particularly in the realm of cancer prediction and diagnostics. These technologies are capable of processing vast amounts of complex data, which is difficult for traditional methods to analyze, allowing for more accurate and timely identification of patients at risk. This approach involves utilizing predictive data modelling methods to forecast the likelihood of an individual's possibility of LungCancer [17]. AI-powered models are able to analyze a wide range of data sources—medical imaging, clinical history, lifestyle behaviors, genetic factors, and environmental exposures—leading to the

identification of patterns and trends that might otherwise go unnoticed.

While AI models in lung cancer prediction have shown promising results, many existing systems rely heavily on clinical and demographic data. The inclusion of additional, relevant data could further enhance the predictive accuracy and the overall effectiveness of these models [18]. One such area with substantial potential is the integration of natural product-based therapeutics into AI prediction models.

Natural products, derived from plants, fungi, and microorganisms, have been extensively studied for their medicinal properties. Many natural compounds are known for their anti-inflammatory, antioxidant, and anticancer effects, and emerging research continues to reveal their potential in preventing cancer progression. These compounds, such as curcumin from turmeric, resveratrol from grapes, and EGCG from green tea, have demonstrated biological activity that could influence cancer development. However, the integration of this knowledge into AI models remains an area that is largely unexplored.

Integrating data on natural product interactions with cancer biology into AI models could transform cancer prevention strategies. The combination of AI's ability to analyze complex datasets with the therapeutic insights from natural products could provide healthcare professionals with more accurate, actionable predictions of lung cancer risk. This, in turn, would allow for more personalized and precise interventions, such as recommending specific natural products, dietary changes, or lifestyle modifications based on an individual's unique risk profile.

The goal of this white paper is to explore the potential of combining AI-driven lung cancer prediction models with the therapeutic properties of natural products. We aim to show how the integration of these two powerful elements—machine learning and natural compounds—can create a more holistic and effective approach to lung cancer prevention. By doing so, we could enable the development of highly personalized, data-driven healthcare solutions that not only predict lung cancer risk with greater accuracy but also offer tangible preventative measures to reduce that risk. In this paper, we will outline the current landscape of AI in lung cancer prediction, highlight the role of natural products in cancer prevention, and propose a framework for integrating these two fields to offer better patient outcomes and lower healthcare costs.

By combining the strengths of both AI technology and natural therapeutics, we can open up new possibilities in lung cancer prevention and management, paving the way for more effective and personalized healthcare solutions in the future [19].

2. BACKGROUND: AI IN LUNG CANCER PREDICTION

Lung cancer remains one of the most critical public health challenges globally, with its high mortality rate being attributed to late-stage diagnoses when the disease is often harder to treat. Early detection of lung cancer has been proven to significantly improve patient outcomes, as it allows for timely interventions and better treatment options. Traditionally, lung cancer detection has relied heavily on imaging technologies such as X-rays and CT scans, alongside clinical assessments of symptoms and risk factors. While these methods have advanced over the years, they are still not foolproof, particularly in the early stages of the disease.

In recent years, artificial intelligence (AI) and machine learning (ML) have shown great potential in enhancing the prediction, detection, and diagnosis of lung cancer [17]. AI algorithms, particularly those based on machine learning, can process and analyze complex datasets far more efficiently than traditional methods. These algorithms are designed to learn from historical data, identifying subtle patterns that may be missed by human experts. This capability makes AI especially valuable in the realm of medical diagnostics, where data volumes are vast and continuously expanding.

Machine Learning Models in Lung Cancer Prediction

Various machine learning techniques have been employed in the development of lung cancer prediction models. Among the most commonly used methods are:

Random Forests (RF): A type of ensemble learning method that combines multiple decision trees to improve prediction accuracy. It has been widely used for classification tasks, including lung cancer detection, due to its robustness in

handling complex, high-dimensional data and its ability to assess the importance of individual variables in predicting outcomes.

Support Vector Machines (SVM): An algorithm that is particularly effective for classification tasks with high-dimensional data. SVMs have been employed in lung cancer prediction to classify patients as at risk or not, based on various features such as age, smoking history, and other clinical markers.

Deep Learning: More recently, deep learning techniques, such as convolutional neural networks (CNNs) and recurrent neural networks (RNNs), have been applied to analyze medical images (e.g., CT scans) and extract patterns that can be indicative of lung cancer. These algorithms are particularly powerful in detecting subtle abnormalities in images, offering an edge in the early detection of tumors that may be missed by human radiologists.

Logistic Regression and Gradient Boosting: These methods have been employed to develop risk prediction models based on clinical and lifestyle data. Logistic regression is useful for determining the probability of an event, such as lung cancer, occurring based on a set of input variables. Gradient boosting algorithms, such as XGBoost, combine multiple weak learners to create a strong predictive model, which has proven effective in predicting outcomes based on complex clinical datasets.

Data Sources for AI-Based Lung Cancer Prediction

For AI models to effectively predict lung cancer risk, they need access to large and diverse datasets. Several types of data are commonly used:

Clinical Data: Patient history, including medical records, family history of cancer, age, gender, and pre-existing conditions, are essential features for developing predictive models. Clinical data also includes lifestyle factors such as smoking habits, exposure to environmental carcinogens, and diet.

Imaging Data: Medical imaging, particularly CT scans and X-rays, provides valuable visual information that can be processed by AI algorithms. Deep learning techniques, such as CNNs, have been shown to be particularly effective in analyzing medical images to detect early-stage tumors and other abnormalities.

Genetic and Molecular Data: Advances in genomics have led to an increase in the availability of genetic data, such as mutations in specific genes that are linked to an increased risk of lung cancer. AI models that integrate genetic information with other types of data could offer more precise predictions.

Environmental and Behavioral Data: Exposure to environmental factors such as air pollution, asbestos, and second-hand smoke has been linked to lung cancer risk. Behavioral data, such as physical activity levels, dietary habits, and alcohol consumption, also contribute to an individual's risk profile.

Challenges in AI-Based Lung Cancer Prediction

While AI offers significant promise in lung cancer prediction, there are several challenges that need to be addressed to fully realize its potential:

Data Quality and Availability: AI models are only as good as the data they are trained on. High-quality, well-labeled datasets are essential for training accurate models. However, access to comprehensive datasets that combine clinical, genetic, imaging, and behavioral data is often limited. Additionally, privacy concerns surrounding patient data make it difficult to create large, publicly accessible datasets.

Model Interpretability: One of the significant challenges with advanced AI models, particularly deep learning models, is their lack of interpretability. While these models may achieve high accuracy, understanding how and why a particular prediction was made remains a challenge. This lack of transparency can hinder trust and adoption among healthcare professionals, who require a clear rationale for the model's decisions.

Integration into Clinical Practice: AI-based models for lung cancer prediction must be integrated seamlessly into clinical workflows for widespread adoption. This requires collaboration between data scientists, healthcare providers, and technology companies to ensure that AI tools are both effective and user-friendly for medical practitioners.

Bias in AI Models: AI models are susceptible to biases in training data, which can lead to inaccurate predictions for certain patient populations. For example, if the training data does not adequately represent certain demographic groups (such as minority populations), the model may produce biased predictions, leading to inequitable healthcare outcomes. Ensuring diversity and representativeness in training datasets is crucial to mitigating this issue.

The Future of AI in Lung Cancer Prediction

Despite these challenges, AI holds immense potential to transform the landscape of lung cancer prediction and treatment. As more data becomes available and machine learning techniques continue to advance, AI models will become increasingly sophisticated, enabling earlier detection and more personalized treatment options. The integration of AI with other emerging technologies, such as genomics and natural language processing (NLP), will further enhance the accuracy of predictions and help clinicians make more informed decisions.

Incorporating data on natural products into these AI models, as explored in this white paper, could lead to even more personalized approaches to lung cancer prevention and treatment. By enhancing traditional AI models with natural therapeutics, healthcare providers could recommend specific lifestyle changes or supplement regimens tailored to an individual's risk profile, offering a more holistic approach to cancer management.

3. NATURAL PRODUCT-BASED THERAPEUTICS AND THEIR ROLE IN LUNG CANCER

Natural products, derived from plants, fungi, and microorganisms, have a long history of use in traditional medicine [20]. In modern scientific research, many natural compounds are being studied for their pharmacological

properties, including anti-cancer effects. These compounds have the potential to impact cancer biology through several mechanisms, such as inhibiting cancer cell proliferation, promoting apoptosis (programmed cell death), suppressing inflammation, and enhancing the immune response.

Lung cancer, one of the deadliest cancers globally, poses significant challenges for treatment, especially when diagnosed at advanced stages. However, the therapeutic potential of natural products offers new avenues for improving both prevention and treatment. By integrating these compounds into a more comprehensive AI-driven lung cancer prediction model, we can explore personalized strategies to reduce cancer risk and optimize treatment approaches.

Mechanisms of Action in Cancer Therapy

Natural products can exert their anticancer effects through various biological mechanisms, such as:

I. Inhibition of Tumor Cell Growth and Proliferation:

Many natural compounds, such as flavonoids and polyphenols, have been shown to inhibit the proliferation of cancer cells. They can interfere with key cellular pathways involved in the cell cycle, thus preventing the uncontrolled growth of tumor cells. By modulating these pathways, natural products help slow down or stop the growth of cancer cells, making them promising candidates for therapeutic development.

II. Induction of Apoptosis:

Apoptosis is a process through which the body eliminates dysfunctional or damaged cells, including cancer cells. Natural compounds like curcumin and resveratrol have demonstrated the ability to induce apoptosis in lung cancer cells. These compounds work by activating specific proteins that signal the cell to self-destruct, thereby reducing tumor burden. By promoting apoptosis in cancer cells, natural products can help control the spread of tumors.

III. Anti-inflammatory Effects:

Chronic inflammation is a recognized factor in cancer development, including lung cancer. Natural products such as curcumin (found in turmeric) and resveratrol (found in grapes) possess potent anti-inflammatory properties that help reduce inflammation in the body. These anti-inflammatory effects can decrease the risk of cancer cell formation, prevent tumor growth, and suppress metastasis (the spread of cancer to other organs). By reducing inflammation, natural products could also improve overall patient health and mitigate the adverse effects of chemotherapy and other conventional treatments.

IV. Inhibition of Angiogenesis:

Tumors require a blood supply to grow and spread, a process known as angiogenesis. Several natural compounds have been identified as anti-angiogenic agents, meaning they can inhibit the formation of new blood vessels that supply tumors. For example, epigallocatechin gallate (EGCG), a polyphenol found in green tea, has shown potential to inhibit angiogenesis in lung cancer by blocking the signaling pathways involved in blood vessel formation. By halting angiogenesis, these compounds help starve the tumor of nutrients and prevent it from growing further.

V. Immune System Modulation:

The immune system plays a critical role in identifying and eliminating cancer cells. Some natural products can enhance the immune response, making it more efficient in recognizing and destroying cancer cells. For instance, polysaccharides extracted from medicinal mushrooms have been shown to activate immune cells like macrophages and dendritic cells, which are involved in the detection and killing of tumor cells. By boosting immune function, natural products can potentially work alongside other therapies to improve treatment outcomes and reduce cancer recurrence.

Key Natural Products and Their Role in Lung Cancer

Several natural products have been extensively researched for their anti-cancer properties, particularly in lung cancer. Below are some of the most promising natural compounds and their roles in combating lung cancer:

I. Curcumin (Turmeric):

Curcumin is a polyphenolic compound derived from turmeric, known for its potent anti-inflammatory, antioxidant, and anticancer properties. Studies have shown that curcumin inhibits the proliferation of lung cancer cells, induces apoptosis, and suppresses metastasis. It does so by modulating several key signaling pathways, including NF-kB and MAPK, that are involved in cell survival and cancer progression. Furthermore, curcumin has shown potential in overcoming chemotherapy resistance, making it a valuable adjunct in lung cancer treatment.

II. Epigallocatechin Gallate (EGCG):

EGCG is a powerful antioxidant and the main active component of green tea. It has been shown to inhibit lung cancer cell proliferation and induce cell death by activating the p53 pathway, a key tumor suppressor gene. EGCG also acts as an anti-angiogenic agent by inhibiting the vascular endothelial growth factor (VEGF) pathway, which is essential for tumor blood vessel formation. Research suggests that EGCG may enhance the effects of chemotherapy and prevent the recurrence of lung cancer.

III. Resveratrol (Grapes and Red Wine):

Resveratrol, a polyphenolic compound found in grapes, berries, and red wine, has garnered attention for its anticancer properties. It works by inhibiting the growth of cancer cells, promoting apoptosis, and reducing inflammation. Resveratrol has shown promise in preclinical studies for lung cancer, where it downregulates the expression of various proteins involved in tumor progression, such as matrix metalloproteinases (MMPs). It may also help protect normal lung cells from the damage caused by chemotherapy and radiation therapy.

IV. Quercetin (Apples, Onions, and Other Fruits):

Quercetin is a flavonoid found in many fruits, vegetables, and grains, and has been studied for its anticancer, anti-inflammatory, and antioxidant properties. In lung cancer, quercetin has shown potential in inhibiting the growth of cancer cells by blocking key pathways such as PI3K/Akt and MAPK, which are involved in tumor cell survival and proliferation. Quercetin also enhances the effects of chemotherapy drugs by sensitizing tumor cells to the cytotoxic effects of these treatments.

V. Sulforaphane (Cruciferous Vegetables):

Sulforaphane, found in cruciferous vegetables like broccoli and Brussels sprouts, is known for its ability to activate the body's antioxidant defenses and detoxification enzymes. In lung cancer models, sulforaphane has been shown to inhibit the growth of cancer cells and induce apoptosis. It works by modulating the Nrf2 pathway, which regulates the expression of genes involved in oxidative stress response and carcinogen metabolism. Sulforaphane may also reduce the risk of lung cancer by enhancing the body's ability to neutralize carcinogens.

VI. Gingerol (Ginger):

Gingerol, the active component in ginger, has been shown to possess anti-inflammatory, antioxidant, and anticancer properties. In lung cancer, gingerol has demonstrated the ability to inhibit the proliferation of cancer cells and induce cell death through several pathways, including the regulation of the AKT/mTOR signaling pathway. Gingerol may also help alleviate chemotherapy-induced nausea and improve overall patient well-being.

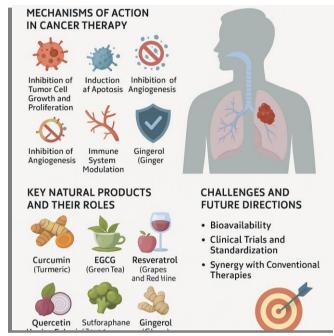


Figure 1. Natural Product-Based Therapeutics in Lung Cancer Treatment: Mechanisms, Key Compounds, and Challenges.

Challenges and Future Directions

Despite the promising anticancer properties of natural products, several challenges remain in their clinical application:

I. Bioavailability:

Many natural compounds are poorly absorbed in the body, limiting their therapeutic potential. Researchers are exploring ways to enhance the bioavailability of these compounds, such as using nanoparticles, liposomes, or co-administration with other agents to improve absorption and delivery.

II. Clinical Trials and Standardization:

Most of the studies on natural products in cancer therapy have been preclinical (in vitro or in animal models), with limited clinical evidence. Further clinical trials are needed to establish their efficacy and safety in humans. Additionally, the standardization of natural products is a significant challenge, as the concentration of active compounds can vary widely depending on the source and preparation.

III. Synergy with Conventional Therapies:

The combination of natural products with conventional cancer treatments like chemotherapy, radiation therapy, or immunotherapy holds great promise. However, careful studies are needed to understand potential interactions, dosage regimens, and the most effective ways to integrate natural products into current treatment protocols.

4. SYNERGY BETWEEN AI MODELS AND NATURAL PRODUCT DATA

The integration of natural product-based data into AI-driven lung cancer prediction models presents a unique opportunity to enhance predictive accuracy and offer personalized, proactive treatment strategies. While AI models, particularly those based on machine learning (ML), have shown significant potential in identifying patterns and predicting cancer risk based on clinical, environmental, and lifestyle data, the inclusion of natural product data could refine these predictions and open new avenues for prevention and treatment. The synergy between AI and natural products leverages the strengths of both fields to create a more holistic and effective approach to lung cancer management

Enhancing AI Prediction Models with Natural Product Data:

AI models are designed to learn from large and diverse datasets, identifying patterns and correlations that may be invisible to the human eye. These models are particularly effective in analyzing clinical data, imaging, and environmental factors to predict the likelihood of lung cancer in individuals. However, the predictive accuracy of these models can be limited by the scope of data they incorporate. Natural product data, which includes information on compounds with potential anti-cancer properties, could significantly enhance these AI models by introducing additional data points that influence tumor development, progression, and response to treatment.

By integrating natural product data, AI models can take into account not just genetic, clinical, and lifestyle factors, but also the therapeutic potential of specific natural compounds that can help reduce cancer risk or enhance treatment efficacy. This enriched dataset would allow the AI to build a more comprehensive and accurate risk profile for individual patients.

A. Personalized Cancer Prevention Strategies

One of the most compelling applications of AI models enhanced with natural product data is the creation of personalized prevention strategies. Today, cancer prevention strategies typically rely on general guidelines based on population-level data. However, these strategies often fail to account for the unique genetic, environmental, and lifestyle factors that contribute to an individual's risk. Natural product data offers an opportunity to personalize these strategies by recommending specific compounds or lifestyle changes based on a patient's unique profile.

For instance, if an AI model detects that a patient has a high genetic predisposition to lung cancer and is also exposed to environmental risk factors like smoking or air pollution, it might recommend a personalized regimen of natural products—such as curcumin, resveratrol, or EGCG—that have demonstrated potential in reducing cancer risk or enhancing detoxification processes. Additionally, dietary and lifestyle changes could be suggested to further mitigate risk. This personalized approach would be more targeted and likely more effective than generic prevention methods, increasing the chances of reducing the onset of lung cancer.

B. Enhancing Treatment Efficacy and Overcoming Drug Resistance

Another significant benefit of integrating natural product data into AI models is the potential to enhance treatment efficacy and overcome drug resistance. Conventional cancer treatments, such as chemotherapy and radiation, often have limited effectiveness due to the development of drug resistance. This resistance occurs when cancer cells adapt to chemotherapy drugs, rendering them less effective over time.

Natural products have shown promise in overcoming this resistance. For example, curcumin has been found to increase the sensitivity of cancer cells to chemotherapy, while EGCG may help reduce resistance to certain chemotherapy drugs. By integrating data on how these compounds interact with chemotherapy agents, AI models can identify potential combinations of treatments that may enhance the efficacy of conventional therapies. AI-driven models could also predict the most effective dosage and timing of natural product supplements to maximize their potential to combat drug resistance.

Furthermore, the integration of natural product data could allow for more precise personalized treatment plans. AI models can analyze individual patient data and recommend the most effective combination of natural products and conventional treatments based on the patient's cancer type, stage, and genetic makeup. This personalized treatment approach could increase treatment success rates and reduce side effects, making therapy more effective and manageable.

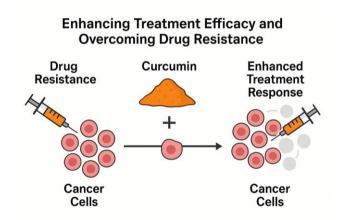


Figure 2. Enhancing Treatment Efficacy and Overcoming Drug Resistance with Curcumin.

C. AI-Driven Natural Product Optimization

AI can also be used to optimize the use of natural products in cancer treatment by identifying new compounds with therapeutic potential. Traditional methods of identifying new natural products often involve a time-consuming and resource-intensive process of screening plants, fungi, and microorganisms for bioactive compounds. However, AI models can accelerate this process by predicting the bioactivity of new compounds based on their chemical structure and known interactions with cancer-related pathways.

For example, AI models can be trained on vast datasets of natural compounds and their known effects on cancer cells, identifying patterns that indicate which compounds are most likely to have anticancer effects. These insights could then be used to prioritize compounds for further testing, saving valuable time and resources in the drug discovery process.

Challenges in Integrating Natural Product Data with AI Models:

Despite the promising potential of integrating natural product data into AI models, several challenges remain:

Data Quality and Availability: High-quality data on the effects of natural products in cancer therapy is often limited, particularly in clinical settings. While preclinical studies (e.g., in vitro and animal studies) show promising results, more human clinical trials are needed to validate these findings. Without robust, high-quality clinical data, AI models may struggle to provide accurate predictions.

Bioavailability of Natural Products: One of the major challenges with using natural products as therapeutics is their bioavailability—the extent to which the body can absorb and utilize the active compounds. Many natural compounds have poor bioavailability, which limits their effectiveness. Advances in drug delivery systems, such as nanoparticles or liposomes, are needed to improve the absorption and distribution of these compounds within the body.

Standardization of Natural Products: Unlike synthetic drugs, natural products can vary significantly in composition depending on their source, preparation, and extraction methods. This variability makes it difficult to standardize their use in clinical settings. AI models must be trained on standardized, high-quality data to make accurate predictions and recommendations.

Ethical and Regulatory Considerations: The use of natural products in cancer treatment is still under investigation, and regulatory approval processes for these compounds are more complex than for conventional pharmaceuticals. Researchers and healthcare providers must ensure that the use of natural products is safe and effective, adhering to rigorous regulatory standards before widespread adoption.

The integration of natural product data into AI-driven lung cancer prediction models holds immense potential for transforming the landscape of cancer prevention, early detection, and treatment. By leveraging AI's ability to analyze complex data alongside the therapeutic properties of natural products, we can create more personalized and effective strategies for managing lung cancer risk. While challenges exist, the potential benefits-ranging from improved early detection to overcoming drug resistance—make this an exciting area of research and innovation in oncology. As research continues to evolve, AI models will become increasingly adept at incorporating diverse datasets, including natural product data, to offer more comprehensive, data-driven healthcare solutions. The future of cancer treatment may lie in the synergy between cutting-edge technologies like AI and the therapeutic potential of natural compounds, offering new hope for patients and improved outcomes across the globe.

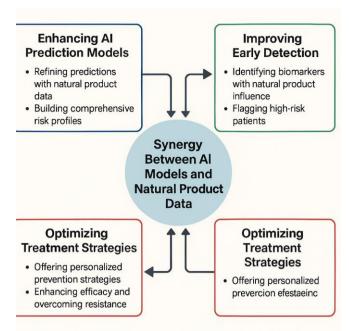


Figure 3. Synergy of AI Models and Natural Product Data in Lung Cancer Prediction and Treatment.

5. THE ROLE OF PERSONALIZED MEDICINE IN CANCER PREVENTION

Personalized medicine represents a paradigm shift in healthcare, moving away from a "one-size-fits-all" approach to a more individualized strategy. It tailors medical treatment and prevention strategies based on a person's genetic, environmental, and lifestyle factors, aiming to provide the most effective care for each unique patient [21]. In the context of cancer prevention, personalized medicine has the potential to drastically improve outcomes by identifying individuals at higher risk, optimizing prevention measures, and offering targeted interventions that align with a patient's unique biology [22].

Lung cancer, in particular, is a prime candidate for personalized medicine because it is influenced by a complex interplay of genetic, environmental, and lifestyle factors. Smoking, environmental toxins, family history, and genetic mutations all contribute to lung cancer risk. Personalized medicine seeks to integrate these factors into a comprehensive strategy that can prevent, detect, and treat lung cancer more effectively. By leveraging advances in genomics, artificial intelligence (AI), and natural product-based therapeutics, personalized medicine offers a more nuanced and effective approach to cancer prevention.

Key Components of Personalized Medicine in Cancer Prevention:

A. Genetic Profiling and Risk Stratification

Genetic profiling involves analyzing a patient's DNA to identify specific genetic mutations or variations that may increase the risk of developing cancer. Certain genetic mutations are strongly associated with an increased risk of lung cancer, such as mutations in the EGFR (epidermal growth factor receptor) gene, which can make tumors more aggressive, or TP53, a tumor suppressor gene. By identifying these genetic markers, doctors can assess a person's genetic predisposition to lung cancer and classify them into different risk categories.

For example, individuals with a family history of lung cancer or who carry high-risk genetic variants may be flagged as high-risk patients. These individuals could benefit from earlier and more frequent screenings, such as low-dose CT scans, as well as personalized lifestyle recommendations aimed at reducing environmental risk factors, such as exposure to carcinogens. Genetic testing may also be used to identify mutations that could be targeted with precision therapies if cancer develops later in life.

B. Tailored Prevention Strategies Based on Lifestyle and Environmental Factors

In addition to genetic factors, personalized medicine takes into account lifestyle choices and environmental exposures, which are major contributors to lung cancer risk. Smoking is the leading cause of lung cancer, but other environmental factors, such as exposure to asbestos, air pollution, and secondhand smoke, also increase risk. Personalized medicine aims to provide tailored prevention strategies based on a person's lifestyle and environmental exposure history.

For example, a smoker who has been diagnosed with chronic obstructive pulmonary disease (COPD) might be flagged for increased lung cancer risk and could benefit from personalized counseling, cessation programs, or specific pharmacological interventions. Additionally, personalized strategies may involve recommending specific natural products, such as EGCG (epigallocatechin gallate) from green tea, which has been shown to have protective effects against cancer, or curcumin, which is known for its anti-inflammatory and anticancer properties.

By integrating genetic data with lifestyle and environmental factors, personalized medicine can offer targeted recommendations to reduce a patient's risk of developing lung cancer, improving long-term health outcomes.

C. Predictive Modeling and Risk Assessment Using AI

AI plays a critical role in personalized medicine by helping to develop predictive models that can assess an individual's risk of lung cancer based on a combination of genetic, lifestyle, and environmental data. Machine learning algorithms, particularly those used in AI-based lung cancer prediction models, analyze large datasets to identify patterns that might not be visible to the human eye. These algorithms are capable of processing complex combinations of genetic markers, clinical history, and environmental exposures to provide a more accurate risk assessment.

By integrating data from genomics, imaging, and clinical history, AI models can predict a patient's likelihood of developing lung cancer with higher precision. For example, AI models can identify subtle interactions between genetic mutations and environmental exposures that might increase cancer risk. This personalized risk profile allows clinicians to offer individualized prevention plans, such as targeted screening programs, lifestyle modifications, or chemoprevention with specific natural products.

D. Pharmacogenomics: Tailoring Preventive Therapies

Pharmacogenomics is an area of personalized medicine that focuses on how a person's genetic makeup affects their response to drugs. In cancer prevention, pharmacogenomic data can be used to identify which patients may benefit most from chemopreventive agents or medications aimed at reducing

cancer risk. Some natural products, such as resveratrol (from grapes) or sulforaphane (from cruciferous vegetables), have demonstrated cancer-preventive properties in laboratory studies.

Pharmacogenomic testing can help determine the best preventive treatment options for individuals based on their genetic profile. For instance, individuals with specific genetic variations may respond better to certain natural products or pharmaceuticals, while others may need different treatments or no intervention at all. This approach ensures that patients receive the most effective, targeted preventive therapies, reducing unnecessary treatments and improving patient outcomes.

E. Natural Product-Based Approaches in Personalized Prevention

Natural products play an increasingly important role in personalized medicine, particularly in cancer prevention. These compounds, derived from plants, fungi, and microorganisms, have been found to possess various biological activities, including anti-inflammatory, antioxidant, and anti-cancer properties. For lung cancer prevention, natural products such as curcumin (turmeric), resveratrol (grapes), and EGCG (green tea) have been extensively studied for their potential to reduce cancer risk and enhance the effects of conventional treatments.

The inclusion of natural products into personalized prevention strategies is promising, especially when tailored to individual genetic profiles. For example, a person with a family history of lung cancer may benefit from specific natural compounds that enhance detoxification pathways or reduce oxidative stress. AI models could play a critical role in identifying the most effective combinations of natural products for different patient profiles, optimizing cancer prevention strategies on an individual basis.

F. Early Detection and Screening in Personalized Medicine

In personalized medicine, the goal is not just prevention, but also early detection. By integrating genetic profiling with advanced imaging technologies and AI-driven prediction models, clinicians can identify individuals who are at high risk of lung cancer and recommend early screening protocols. This is particularly important for high-risk populations, such as smokers, individuals with a family history of lung cancer, and those with certain genetic mutations.

Low-dose CT scans, which are used to detect early-stage lung cancer in high-risk patients, could be incorporated into personalized prevention programs, ensuring that individuals at higher risk undergo more frequent and tailored screenings. Personalized screening schedules, along with specific biomarkers linked to cancer risk, could lead to earlier detection of lung cancer, increasing the chances of successful treatment and better outcomes.

The Future of Personalized Medicine in Lung Cancer Prevention:

The future of personalized medicine in lung cancer prevention lies in the integration of multiple technologies and data sources, including genomics, AI, natural product therapeutics, and lifestyle factors. As our understanding of cancer biology and the human genome expands, personalized prevention strategies will become more sophisticated, offering more precise and effective ways to prevent lung cancer.

Personalized medicine has the potential to revolutionize cancer prevention, offering highly targeted approaches that maximize the effectiveness of prevention and early detection while minimizing unnecessary interventions. By tailoring prevention strategies to the unique genetic and environmental profile of each patient, personalized medicine will enable better outcomes, reduce cancer incidence, and improve the quality of life for individuals at risk of lung cancer. Personalized medicine offers an unprecedented opportunity to advance cancer prevention, particularly for complex cancers like lung cancer. By integrating genetic, environmental, and lifestyle factors, alongside the emerging role of natural product-based therapeutics, personalized medicine can provide highly individualized strategies to reduce lung cancer risk. With the help of AI and advances in pharmacogenomics, these strategies can be further optimized, leading to better prevention outcomes and more effective, targeted treatments for patients.

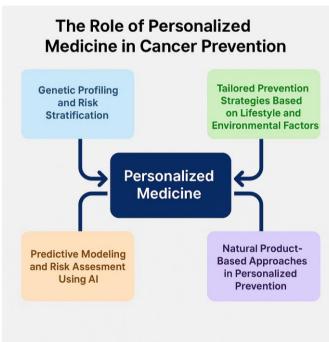


Figure 4. Framework for Personalized Medicine in Cancer Prevention.

6. CASE STUDIES AND EVIDENCE OF NATURAL PRODUCTS IN LUNG CANCER PREVENTION

Natural products have been extensively studied for their potential to prevent, treat, and even reverse the progression of various types of cancer, including lung cancer. Lung cancer, often diagnosed at advanced stages, has a poor prognosis, making prevention and early detection crucial. Several natural compounds derived from plants, fungi, and other organisms have demonstrated promising anticancer properties in laboratory and clinical studies, particularly for lung cancer prevention. This section explores case studies and the scientific evidence supporting the role of natural products in reducing the risk of lung cancer, inhibiting tumor growth, and enhancing the effects of conventional therapies. Traditional methods of tracking progress rely heavily on self-reporting and clinical evaluations, which can be time-consuming and inaccessible to many individuals [23].

A. Curcumin (Turmeric) in Lung Cancer Prevention

Curcumin, the active compound in turmeric, is widely known for its anti-inflammatory, antioxidant, and anticancer properties. Several studies have shown that curcumin can effectively inhibit the growth of lung cancer cells, induce apoptosis (programmed cell death), and reduce metastasis. It achieves these effects by modulating multiple signaling pathways involved in cancer progression, including the NF-kB and MAPK pathways, which are known to regulate inflammation and cell survival.

Case Study: A study published in Clinical Cancer Research (2011) investigated the effects of curcumin on human non-small cell lung cancer (NSCLC) cells. The study found that curcumin significantly inhibited the proliferation of lung cancer cells and induced apoptosis through the p53 pathway, a tumor suppressor gene that regulates cell cycle arrest and apoptosis. Additionally, the study demonstrated that curcumin reduced the invasive capacity of lung cancer cells, suggesting its potential as a chemopreventive agent.

Further studies have indicated that curcumin can work synergistically with chemotherapy agents, such as paclitaxel and cisplatin, by enhancing their efficacy and overcoming drug resistance. This makes curcumin a potential adjunct to standard cancer therapies, improving patient outcomes without significant toxicity.

B. Epigallocatechin Gallate (EGCG) from Green Tea

EGCG is a polyphenol found in green tea that has been extensively studied for its anticancer properties. It acts as a potent antioxidant, preventing oxidative stress, and has been shown to inhibit cancer cell proliferation, angiogenesis (the formation of new blood vessels), and metastasis. EGCG also modulates key molecular pathways involved in cancer progression, such as AKT/mTOR, Wnt/ β -catenin, and MAPK/ERK.

Case Study: A clinical trial published in Cancer Prevention Research (2013) explored the effects of EGCG on lung cancer patients at high risk [24]. The study involved administering EGCG to individuals with a history of heavy smoking and observed significant reductions in biomarkers associated with cancer cell growth and inflammation. The study concluded that EGCG supplementation could potentially reduce the risk of lung cancer by inhibiting tumor-promoting pathways.

EGCG has also been found to sensitize cancer cells to radiation therapy and chemotherapy. For instance, in preclinical studies, EGCG enhanced the effects of radiotherapy in lung cancer models by increasing cancer cell apoptosis and reducing tumor growth. This suggests that EGCG, as a natural adjunct to conventional treatments, could play a crucial role in improving lung cancer outcomes.

C. Resveratrol from Grapes and Red Wine

Resveratrol is a polyphenolic compound found in grapes, red wine, and certain berries. It has been shown to possess antioxidant, anti-inflammatory, and anticancer properties. Resveratrol works by inhibiting cell cycle progression, inducing apoptosis, and blocking angiogenesis in cancer cells. It also inhibits the activity of matrix metalloproteinases (MMPs), enzymes involved in tumor invasion and metastasis.

Case Study: A study published in the Journal of Nutritional Biochemistry (2010) investigated the effects of resveratrol on lung cancer prevention in animal models. The study found that resveratrol significantly reduced the number of lung cancer nodules and inhibited the progression of tumors. It was concluded that resveratrol could be a valuable agent in the prevention of lung cancer, especially for individuals exposed to environmental carcinogens like cigarette smoke.

Clinical studies have also demonstrated the potential of resveratrol in overcoming chemotherapy resistance. In a study published in Molecular Cancer Therapeutics (2014), resveratrol was shown to enhance the effects of cisplatin, a commonly used chemotherapy drug, by modulating the PI3K/Akt signaling pathway and increasing cancer cell sensitivity to the drug. This suggests that resveratrol could serve as a complementary agent in lung cancer therapy.

D. Quercetin from Apples, Onions, and Other Fruits

Quercetin is a flavonoid found in a variety of fruits, vegetables, and grains, including apples, onions, and berries. Quercetin has been shown to possess anticancer, anti-inflammatory, and antioxidant properties. It exerts its anticancer effects by inhibiting cell proliferation, inducing apoptosis, and suppressing angiogenesis. Quercetin also modulates multiple signaling pathways, including the PI3K/Akt and MAPK/ERK pathways, which play a key role in cancer cell survival and growth.

Case Study: A preclinical study published in Cancer Letters (2012) explored the effects of quercetin on lung cancer cells. The study found that quercetin significantly inhibited the proliferation of lung cancer cells and induced cell cycle arrest at the G1/S phase. Moreover, quercetin reduced the expression of cyclin D1, a protein involved in cell cycle regulation, suggesting its potential as a therapeutic agent for lung cancer prevention.

Another study published in The Journal of Nutritional Biochemistry (2011) demonstrated that quercetin could enhance the effectiveness of chemotherapy by reducing drug resistance in lung cancer cells. The study showed that quercetin, when combined with docetaxel (a chemotherapy drug), increased the sensitivity of lung cancer cells to the drug, potentially improving the outcomes of chemotherapy treatments.

E. Sulforaphane from Cruciferous Vegetables

Sulforaphane is a bioactive compound found in cruciferous vegetables like broccoli, Brussels sprouts, and cabbage. It has potent anticancer properties, primarily through its ability to activate the Nrf2 pathway, which regulates the body's antioxidant defenses and detoxification enzymes. Sulforaphane has been shown to inhibit cancer cell proliferation, induce apoptosis, and block angiogenesis.

Case Study: A study published in The Journal of Clinical Investigation (2008) investigated the effects of sulforaphane on lung cancer cells [26]. The study found that sulforaphane effectively inhibited the growth of lung cancer cells and induced apoptosis by upregulating the expression of p21 and p53, tumor suppressor proteins involved in cell cycle regulation and apoptosis. Furthermore, sulforaphane enhanced the ability of lung cancer cells to detoxify carcinogens, reducing their potential for DNA damage and mutation.

Sulforaphane has also been shown to work synergistically with other natural compounds, such as EGCG, to enhance their anticancer effects. In a study published in Cancer Research (2013), a combination of sulforaphane and EGCG was found to significantly reduce the proliferation of lung cancer cells and suppress tumor growth in animal models.

F. Gingerol from Ginger

Gingerol is the active compound in ginger, known for its anti-inflammatory, antioxidant, and anticancer effects. Gingerol has been shown to inhibit lung cancer cell proliferation and reduce the inflammatory responses that contribute to tumor growth. It also modulates several molecular pathways involved in cancer progression, including the NF-kB and MAPK pathways.

Case Study: A study published in The Journal of Alternative and Complementary Medicine (2010) investigated the effects of gingerol on lung cancer prevention. The study found that gingerol significantly inhibited the growth of lung cancer cells and induced apoptosis. The study also observed that gingerol reduced the expression of inflammatory cytokines, suggesting its potential in reducing lung cancer risk, particularly in individuals with chronic inflammation or respiratory conditions.

The case studies and evidence presented demonstrate the promising role of natural products in lung cancer prevention. Compounds like curcumin, EGCG, resveratrol, quercetin, sulforaphane, and gingerol have shown considerable potential in inhibiting lung cancer cell proliferation, inducing apoptosis, and preventing metastasis. Moreover, natural products have demonstrated the ability to work synergistically with conventional cancer therapies, enhancing their effectiveness and overcoming drug resistance.

While many of these findings come from preclinical studies, clinical trials are essential to further validate the safety and efficacy of natural products in lung cancer prevention and treatment. Ongoing research will help establish these natural compounds as valuable adjuncts to current treatment strategies, offering new hope for lung cancer patients and individuals at high risk.



Figure 4. Natural Products in Lung Cancer Prevention: Key Compounds and Their Mechanisms.

7. PROPOSED FRAMEWORK FOR AI AND NATURAL PRODUCT INTEGRATION

The integration of artificial intelligence (AI) with natural product-based therapeutics offers a novel approach to lung cancer prediction and treatment. While AI has revolutionized healthcare by enabling more accurate and efficient predictions, its effectiveness can be further enhanced by incorporating data on natural compounds that have proven anticancer properties. The proposed framework for integrating AI with natural product data involves several key steps, from data collection and integration to personalized treatment recommendations.

This framework aims to create a holistic approach to lung cancer management, where AI models predict risk and recommend personalized strategies while leveraging the therapeutic potential of natural products. Below is a detailed step-by-step guide to this integration:

1. Data Collection

Clinical Data: Includes patient medical records, genetic information, lifestyle factors, and imaging.

Natural Product Data: Includes information on natural compounds like curcumin, EGCG, and resveratrol, focusing on their mechanisms, dosages, and safety profiles.

Environmental Data: Factors like smoking history and exposure to carcinogens are also considered.

2. Data Integration

Feature Engineering: Identifying relevant features from both clinical and natural product data.

Data Fusion: Merging diverse data sources into a unified model to create a comprehensive risk profile.

Multi-Modal Learning: Combining clinical, genetic, and natural product data into one AI model for more accurate predictions.

3. Model Training and Testing

Supervised Learning: AI models are trained using labeled data to predict cancer risk and personalize treatment plans.

Unsupervised Learning: AI models detect hidden patterns that could inform prevention or treatment strategies.

Cross-Validation: Ensuring the model's accuracy and robustness through multiple tests.

4. Personalized Prevention and Treatment Recommendations

Risk Prediction: AI models predict lung cancer risk based on individual profiles.

Personalized Prevention: AI suggests targeted lifestyle changes and natural product-based interventions (e.g., curcumin, EGCG).

Treatment Plans: For patients diagnosed with lung cancer, AI recommends personalized treatment options, combining

conventional therapies with natural products to improve efficacy.

5. Future Considerations

Clinical Integration: Ensuring AI models are seamlessly integrated into clinical workflows.

Data Privacy: Protecting patient data and adhering to privacy regulations (e.g., HIPAA).

8. CONCLUSION

This white paper has delved into the promising synergy between natural product-based therapeutics and AI-driven lung cancer prediction models, showcasing the potential for a more comprehensive and personalized approach to cancer prevention, early detection, and treatment. By combining the cutting-edge capabilities of artificial intelligence with the proven therapeutic properties of natural compounds, this interdisciplinary approach could revolutionize the way we predict, prevent, and manage lung cancer.

AI has already demonstrated its power in processing complex datasets to predict lung cancer risk and guide treatment decisions. Machine learning models, trained on large-scale clinical, genetic, and environmental data, have proven to be effective in identifying high-risk individuals and offering early detection strategies. However, these models often rely on clinical and demographic data alone, which limits their ability to offer personalized prevention strategies tailored to individual biological profiles.

Integrating natural product data into AI models enhances their predictive capabilities by introducing an additional layer of therapeutic insight. Natural products such as curcumin, EGCG, resveratrol, and others have shown significant potential in preventing cancer progression, inhibiting tumor growth, and sensitizing cancer cells to traditional therapies. These compounds, derived from everyday foods and plants, are widely accessible and may offer cost-effective, low-risk interventions for individuals at high risk of developing lung cancer. By leveraging this data, AI models can recommend personalized natural product-based interventions, lifestyle changes, and dietary modifications that specifically address the unique risk factors of each patient.

The combination of AI's analytical power and the diverse mechanisms of action found in natural products creates a robust framework for personalized medicine in lung cancer prevention.

In conclusion, the integration of natural product-based therapeutics with AI-driven lung cancer prediction models holds tremendous potential to redefine the way we approach cancer prevention, early detection, and treatment. By leveraging the strengths of both AI and natural products, we can create more accurate, personalized strategies that ultimately improve patient outcomes, reduce healthcare costs, and lower the global burden of lung cancer. This interdisciplinary approach marks a significant step forward in the ongoing fight against cancer, bringing us closer to more effective and accessible solutions for cancer prevention and treatment.

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