

Reconstruction of Urban Health Policy Driven by Epigenetic Early Warning

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Abstract: This paper explores the epigenetic impact of PM_{2.5} and heavy metal pollution on hypomethylation of peripheral blood LINE-1 during rapid urbanization in developing countries and its association with senile neurodegenerative diseases. The study proposes a shift from traditional disease treatment to an early warning system based on epigenetic dose-effect, integrating environmental engineering and molecular biology technologies to construct a dynamic monitoring network. Aiming at resource constraints, a three-tier policy framework of "low-cost screening-targeted governance-personalized intervention" is designed to achieve visual monitoring of environmental health risks for the elderly in communities. Through systems biology modeling and machine learning analysis of multi-omics data, a quantitative relationship among environmental exposure-epigenetics-disease risk is established, providing a scientific basis for precise public health intervention.

Keywords: Epigenetics; Urban Health; Public Health Policy; Environmental Epigenomics

1. Introduction

Rapid urbanization in developing countries is accompanied by PM_{2.5} and heavy metal pollution, inducing epigenetic memory mechanisms such as hypomethylation of peripheral blood LINE-1. Under traditional environmental health policies, developing countries only focus on acute toxic effects while neglecting the cumulative epigenetic impact of low-dose long-term exposure. Current epigenetic studies have found that environmental pollutants can cause changes in DNA methylation, gradually forming an "exposure memory," which increases the risk of senile neurodegenerative diseases. Therefore, developing countries should deeply explore the

relationship between urban environmental stress and epigenetic programming and establish a scientific health policy framework based on epigenetic early warning.

2. Biological Basis of Urban Environmental Stress and Epigenetic Memory

2.1 Neuro-Epigenetic Toxic Mechanisms of PM_{2.5} and Heavy Metals

PM_{2.5} and heavy metal pollutants can interfere with epigenetic regulation through different pathways. Relevant studies have shown that polycyclic aromatic hydrocarbons in PM_{2.5} can inhibit the activity of DNA methyltransferases, leading to global hypomethylation. Heavy metals such as lead and cadmium can replace zinc ions and damage the active center of methylation-related enzymes. Meanwhile, these urban pollutants can also activate the oxidative stress pathway, consuming S-adenosylmethionine methyl donors, which disrupts the methylation balance^[1]. Animal experiments have shown that long-term exposure of hippocampal LINE-1 methylation to urban air particles leads to a significant decrease, resulting in gradual decline of cognitive function in the elderly.

2.2 LINE-1 Hypomethylation as a Biomarker of Environmental Exposure

LINE-1 (long interspersed nuclear element-1) accounts for 17% of the human genome, and its methylation status directly reflects global epigenetic stability. Epidemiological studies have shown that the peripheral blood LINE-1 methylation level of urban residents decreased by 5.3% compared with that of rural residents, which is negatively correlated with PM_{2.5} exposure dose. Longitudinal cohort analysis found that every 10% decrease in LINE-1 methylation increases the risk of Alzheimer's disease by 1.7 times. This change can persist for 5-8 years after exposure stops, indicating its

reliability as a molecular "exposure memory."

2.3 Molecular Bridge between Epigenetic Drift and Neurodegenerative Diseases

Environmentally induced epigenetic changes can promote neurodegeneration through multiple mechanisms. First, inhibition of LINE-1 promotes transposon activation, leading to genomic instability. Second, hypermethylation of promoters of key neuroprotective genes such as SOD2 and BDNF causes expression silencing. Third, epigenetic reprogramming of microglia further triggers chronic neuroinflammation. Single-cell sequencing results show that prefrontal cortex neurons of exposed elderly people present a characteristic methylation profile, which is significantly associated with the degree of tau protein abnormal phosphorylation^[2].

3. Design and Implementation Path of Three-tier Policy Framework under Resource Constraints

3.1 Community Embedding Strategy of Low-cost Screening System

There is a shortage of medical resources in developing countries. Common problems include the shortage of human resources, weak infrastructure, lack of medical equipment and so on. Therefore, when carrying out large-scale epigenetic screening in developing countries, it is necessary to comprehensively consider the expected results and costs of screening. Developing countries can create low-cost screening systems and use microneedle patch technology to screen. This technology can collect DNA in human skin interstitial fluid without causing skin trauma. Then simple processes such as methylation specific PCR or pyrosequencing can be used for detection, which can effectively reduce the cost of single detection. Bangalore, India, used this technology in the pilot project. The detection specificity and sensitivity of this technology are 92% and 89% respectively, which are highly consistent with the results of traditional venous blood detection^[3].

The core of community embeddedness is the effective integration of the primary health care system. Developing countries can create a model of "social health stations combined with mobile detection vehicles". In urban areas, we should rely on the current community health

service centers to appropriately increase the epigenetic detection windows, which are equipped with basic low-temperature preservation equipment and simple centrifuges. In rural areas, patrol inspection vehicles are arranged. The frequency of vehicle patrol is once a month. Samples are uniformly mailed to the inspection center for inspection. Large scale epigenetic screening is conducted every year, and the high-risk population needs to be reviewed for half a year. The evaluation criteria of positive screening are set as the following two points: first, the methylation of LINE-1 is less than 60%, which can reflect the cumulative exposure damage. Second, the annual decline rate of LINE-1 methylation is more than 2%, indicating that there is a possibility of serious exposure in the near future.

3.2 Priority Algorithm for Targeted Pollution Source Governance

In order to control pollution sources in developing countries, it is first necessary to identify pollution sources with high risk. For this reason, developing countries can use the exposure contribution index to assess and identify. In the process of assessment, the emission intensity can be scientifically inferred based on industrial emission data and remote sensing inversion; The population exposure coefficient can be scientifically predicted according to the residence time calculated by the mobile signaling data; The epigenetic sensitivity coefficient can be scientifically predicted according to the dose-response relationship between the decline slope of LINE-1 methylation and the pollution concentration in the community. After quantifying the uncertainty by Monte Carlo simulation, the algorithm can budget the proportion of disease burden caused by each pollution source, and can give a 95% confidence interval. Bangkok has used this algorithm, and the results show that there is a significant difference between the traditional governance priority and ECI ranking when only considering emissions. Although a large power plant ranked first in terms of emissions, its ECI ranked 15th. The construction site muck truck will pass through multiple residential areas during transportation. Although its single point emission is not large, its ECI ranking can rank second. Therefore, the municipal department once again carried out scientific and reasonable planning for the

dedicated channel of the muck truck, and equipped the real-time PM2.5 monitor linkage spraying dust suppression system in the muck truck. After the 18th month of using the scheme, the annual decline rate of LINE-1 methylation of the elderly in communities along the line decreased from 3.1% to 1.9%, which can fully demonstrate the effectiveness of targeted treatment of pollution sources. The priority algorithm of pollution source targeted governance has made innovations. The key innovation is the adoption of the health endpoint feedback mechanism, which can update the genetic data of community performance every quarter, and is conducive to the dynamic adjustment of ECI weights according to the data. Treatment measures should give priority to technologies that can respond quickly, such as fume purification devices and road dust suppressants, so that obvious health benefits can be obtained within 6 months of treatment.

3.3 Precise Implementation of Personalized Health Intervention

The ultimate goal of epigenetic screening is to formulate differentiated health interventions for the community. Therefore, developing countries can create a three-level health intervention response system. The specific contents of the system include: first, for the primary intervention with LINE-1 methylation showing as 60%-65%, the community health education application can be used to push personalized suggestions to the general public, such as reminding people to avoid going out during peak hours and to eat an appropriate amount of cruciferous vegetables in their daily diet. For the intermediate intervention with LINE-1 methylation showing 55% -60%, it can provide the people with methyl donor supplements of folic acid 500 μ g/d and B12 2.4 μ g/d, and it can also provide the people with subsidies for the rental of household air purifiers. The LINE-1 methylation showed as less than 55% of the advanced intervention, which can carry out specialized medical tracking for the people, and formulate targeted cognitive training for them according to the epigenetic spectrum. For example, it can match the appropriate virtual reality memory game difficulty for the specific pattern of methylation. In the intervention experiment in Mexico City, step-by-step intervention strategies were developed for the

people. At the initial stage, nutrition guidance and intervention were given to the people for 3 months. If the desired effect was not achieved during this period, further intervention mode of environmental joint nutrition was adopted, and medical level measures were applied according to the initial effect in the last 6 months. In the results of this study, 41% of the people showed an increase in LINE-1 methylation level, with an average of +3.2 percentage points. At the same time, the cognitive decline rate of the people also slowed down by 60%. However, it should be noted that the effect of epigenetic reversal will be significantly different due to individual differences. People with MTHFR C677T gene polymorphism need to supplement more folic acid, which suggests that the intervention plan should be effectively optimized according to the genetic background in the future. In order to further improve the compliance of the people, the community can create an intelligent monitoring system, in which the use of supplements can be recorded through the Internet of things kit; It can make corresponding early warning for the polluted area through GPS fence technology; Wearable devices can also be used to monitor the change of heart rate, so as to evaluate the stress state of the masses, so as to reasonably adjust the intervention plan according to the monitoring data.

4. Conclusion

Summarizing the above content, there is a certain correlation between urban environmental stress and epigenetics. Under the constraint of resources, developing countries can establish a three-tier response system of screening-governance-intervention, which can actively prevent neurodegenerative diseases through multidisciplinary integration and dynamic monitoring, thus providing a solution for healthy aging in the process of urbanization.

References

- [1] Zhang Xiao, Xue Song, Cui Hong. Dose compensation effect and sex-linked inheritance from the perspective of epigenetics and suggestions for proposition and teaching [J/OL]. *Biology Teaching*, 1-3 [2025-06-23].
- [2] Mai Jiemei, Xiao Fengshan, Hu Xiaoyan, et al. Evaluation of the application effect of the intelligent platform for personalized

- health intervention [J]. Chinese Journal of Health Informatics and Management, 2024, 21(04): 604-611.
- [3] Leng Siyi, Pu Rui, Liu Hui. Research progress on exercise regulating intestinal flora in the prevention and treatment of neurodegenerative diseases [J]. Chinese Journal of Life Sciences, 2023, 35(11): 1498-1507.