



Path Analysis: Public Health Impacts Overview in High Natural Background Radiation Area in Botteng, West Sulawesi, Indonesia

Elanda Fikri^{1,2*}, Evan Puspitasari³, Amar Sharaf Eldin Khair⁴

¹Department of Environmental Health, Bandung Health Polytechnic, North Cimahi, Indonesia, 40514

²Center of Excellence on Utilization of Local Material for Health Improvement

³Center For Technology of Radiation Safety and Metrology, Jalan Lebak Bulus No. 49, Jakarta Selatan, Indonesia

⁴Omdurman Islamic University, Geography Department, Omdurman City, Sudan

*Corresponding Author. Email : elandafikri@yahoo.com

Abstract

Introduction: Ionizing radiation exposure affect human being by chromosome aberrations. Botteng has the highest annual radiation exposure rate in Indonesia (6.15 ± 0.81 mSv/year), thus the inhabitants were exposed to continuous low dose radiation. This study aimed to describe the radiation exposure pathway to inhabitants and the cytogenetic response in the high natural background radiation area.

Methods : This cross-sectional study was conducted on 61 residents in Botteng, who were chosen randomly from 9 different exposure rate areas. The path analysis model was used to determine the linear relationship between internal and external radiation dose to the frequency of chromosomal aberrations in lymphocyte cells.

Results: This study found that the external and internal radiation dose rate was 5.49 mSv/year and 10.34 mSv/year, respectively. The effective dose rate of lymphocyte cells was 1.92 mSv/year, and the mean of chromosome aberrations frequency was 0.00082 of approximately 14.695 metaphase cells observed. There was no correlation between the external dose and the chromosome aberrations frequency. There was a linear relationship between the internal dose and the chromosome aberrations frequency ($F=6.634$, $p=0.013$). The internal and external radiation dose simultaneously affects the effective dose ($R^2=0.901$, $p=0.0001$), The Internal and external radiation doses affect the chromosome aberrations frequency of peripheral blood lymphocytes only through effective doses ($R^2=0.093$, $p=0.017$).

Conclusion: The effect of ionizing radiation exposure to chromosome aberrations was found on the inhabitants of high natural background radiation area in Botteng, West Sulawesi. Public health measures and further study should be conducted.

Keywords: chromosome aberrations, natural background radiation, lymphocyte, public health, radio-adaptive response

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Introduction

Exposure of ionizing radiation sources from nature to humans occurs continuously and is inevitable for every living being on earth. The estimated

average annual exposure from natural background radiation sources is 2.4 mSv with a range of 1-10 mSv/year. The natural background radiation sources in question consist of external exposure from cosmic

radiation (0.4 mSv/year) and terrestrial gamma radiation (0.5 mSv/year), as well as internal exposure, namely inhalation (1.3 mSv/year, especially Radon) and ingestion (0.3 mSv/year).¹

Exposure to ionizing radiation comes from cosmic rays and radionuclides in soil, indoor and outdoor air, water, food chain, and so on leads to chronic exposure originating from external and internal radiation. Those chronic exposures resulted in an annual effective dose for the inhabitants. An area with several chronic exposures that causes inhabitants' effective dose to be above the specified level is known as High Natural Background Radiation (HNBR) areas.²

The average data of environmental gamma radiation dose rate in Indonesia is 68 nSv/hour. Areas with a dose rate above the average rate were North Sumatra 186.08 ± 3.72 nSv/hour, West Kalimantan 349.48 ± 57.21 nSv/hour, and West Sulawesi 487±103 nSv/hour. In the results of the 2013 study in West Sulawesi, a dose rate of 272 ± 3 nSv/hour was obtained.³

Mamuju Regency has the highest dose rate in West Sulawesi. Measurements of the external radiation level are taken on radiation source from outside the human body. The average level of exposure to gamma radiation in Mamuju Regency is 631.4 ± 569.5 nSv/hour. The results of a follow-up survey conducted in several villages in Mamuju Regency showed that Botteng Village, Simboro District, is the village with the highest gamma radiation level, namely 700 – 1000 nSv/hour with an annual average of 6.15 ± 0.81 mSv.⁴

Ionizing radiation exposure that occurred in Botteng Village, Mamuju Regency is exposed by low dose chronic radiation exposure. Our Understanding evolved from “small dose may cause harm” to “small dose definitely will cause harm”. Every ionizing exposure causes DNA damage, which is detrimental to health and leads to higher risks of genetic abnormalities. In high natural background radiation area, inhabitants need multidisciplinary of science observations continuously, for supporting inhabitant's health. The researcher was interested in conducting an epidemiological study of environmental health, using the principle of node theory. This cross-sectional study covered environment epidemiological review on the relationship between natural ionizing radiation exposure and the detriments on inhabitant's health. The Pathway model is presented here to demonstrate the direct and indirect mechanism relations between natural ionizing radiation exposure to the detriment of inhabitant health. The pathway is expected to provide recommendations for further public health research.

Methods

The research method used the dynamics principle of environmental epidemiological, namely the node theory (**Figure 1**). The research used a cross-sectional research design, meaning that the research variables (external radiation dose, internal radiation dose, effective dose, and Chromosome Aberrations Frequency) were collected simultaneously.

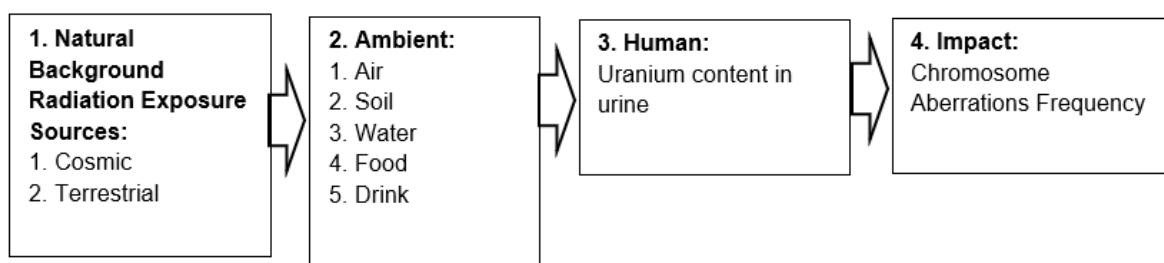


Figure 1. The environmental epidemiological dynamics principle

The study was conducted by processing and analyzing secondary data from a survey conducted by PTKMR – BATAN. Data processing and analysis were carried out in August. The total population of the study was 61 subjects, selected randomly, from 9 hamlets in Botteng Village, Mamuju Regency.

Cosmic and terrestrial natural radiation sources contained in the air, soil, water, food, and drink expose the inhabitants by means of internal exposure and external exposure. The calculation of the external dose is carried out by installing a survey meter, namely the Exploranium GR 135 device. The calculation of the internal dose is carried out by taking samples of food, soil, water, and air for further measurement of the internal radiation dose using a Gamma Spectrometer, while the calculation of the chromosome aberrations frequency is carried out by taking peripheral blood samples and observing the chromosome aberrations frequency using the staining method of Fluorescence In Situ Hybridization (FISH) by Whole Chromosome Probe (WCP) labeled with Fluorescent Isothiocyanate (FITC).

The obtained secondary data of external and internal radiation doses are then processed to determine the effective dose using the following equation.⁵

$$E_T = H_p(10) + \sum_j e(g)_{j,ing} I_{j,ing} + \sum_j e(g)_{i,inh} I_{j,inh}$$

ET = Effective Dose (Sv).

H_p(10) = Equivalent dose of radiation penetration at a depth of 10 mm obtained from individual dosimetry readings (Sv).

e(g)_{j,ing} = Effective dose bound per unit input through digestion for radionuclide J by age group g as stated in the attachment of BAPETEN Regulation No. 4 of 2013 (Sv.Bq-1).

e(g)_{j,inh} = Effective dose bound per unit of intake through inhalation for radionuclide J by age group g as stated in the attachment of

BAPETEN Regulation No. 4 of 2013 (Sv.Bq-1).

I_{j,ing} = Digestive input of radionuclide J (Bq).

I_{j,inh} = Inhalation of radionuclide J (Bq).

The path analysis model is used to determine the linear relationship between internal and external radiation dose to the frequency of chromosomal aberrations in lymphocyte cells. The pattern of the large relationship between internal and external exposure doses directly and indirectly to the frequency of chromosomal aberrations in lymphocyte cells, through effective doses as moderator variables, inhabitant in areas with high natural radiation background. To find out the direct and indirect relationship, it was tested in two structures of path analysis.

Results

Levels of Internal Radiation Dose, External Radiation Dose, and Effective Dose and Chromosome Aberrations Frequency in Lymphocyte Cells

The area of West Sulawesi, Mamuju Regency, Botteng Village, covers 9 Hamlets. Data from univariate analysis are presented in **Table 1** with a central tendency (mean value). The univariate analysis results showed that the average dose of external and internal radiation received by the inhabitants of Botteng Village, West Sulawesi was higher than the world annual dose reported by UNSCEAR.¹

The chromosome aberrations frequency of peripheral blood lymphocytes for 61 residents of Botteng Village, has an average value of 0.00082 from approximately 14,695 metaphase cells observed manually with a microscope. The lowest chromosome aberrations frequency of peripheral blood lymphocytes was 0 and the highest was 2 with a median value of 0.

The residents of Botteng Village experienced the highest chromosome aberrations frequency of peripheral blood lymphocyte of 1.80 mSv/year.

Table 1. Overview of External, Internal, Effective Radiation Doses, and Chromosome Aberrations Frequency in Peripheral Blood Lymphocyte Cells

Variables	N	Mean	Median	Minimum - Maximum	95 % CI
External Dose	61	5.488	5.370	2.630 – 14.590	5.102 – 5.873
Internal Dose	61	10.340	10.120	1.230 – 20.61	9.417 – 11.262
Effective Dose	61	1.918	1.860	1.130 – 3.230	1.801 – 2.035
Chromosome Aberrations Frequency per cell	14.695	0.00082	0.000	0 – 0.2	0.000 – 0.002

The examination results of chromosome aberrations in peripheral blood lymphocyte cells of inhabitants in Botteng Village, West Sulawesi, showed that there were no chromosomal abnormalities in the form of dicentric or acentric fragments, as well as rings. Only a few cells contain 1-4 dicentrics.

Table 2 and **Figure 2** show the overall simultaneous hypothesis test result of structure 1. The F value was 262.483 with a p-value of 0.000, which means that H_0 was rejected. Thus, the external (X_1) and internal (X_2) radiation dose had a simultaneous and significant effect on the

effective dose (Y). The coefficient of determination for structural 1 was $R^2 = 0.901$, then the magnitude effect of other variables outside the model (error) on the effective variable dose (Y) was $1 - R^2 = 0.315$. This means that 90.1% of the Effective dose (Y) can be predicted by the external radiation dose (X_1) and the internal radiation dose (X_2). The other 9.9% was affected by other variables outside the model. So it can be concluded that the effective dose received by lymphocyte cells was strongly affected by the amount of external and internal radiation dose.

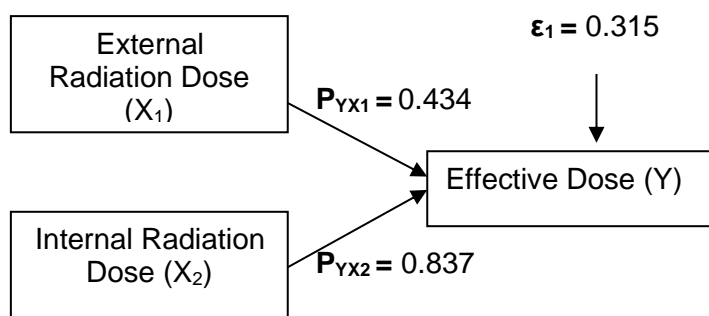


Figure 2. Structure 1 Empirical Causal Relationship of X_1 and X_2 variable towards Y

Table 2. Overall Simultaneous Hypothesis Test Results of Structure 1

	R	R^2	$\frac{\epsilon_1}{\sqrt{1-R^2}}$	F	Standard Error	p
Model 1	0.949	0.901	0.315	262.483	0.14636	0.000

Table 3 and **Figure 3** show the overall simultaneous hypothesis test results of structure 2 with a p-value of 0.87, therefore H_0 is accepted. Model 2 and model 3 have p-value of 0.039 and 0.017, respectively. The coefficient of determination for structural 2 is $R^2 = 0.093$, therefore the magnitude of other variables

effects outside the model (error) towards the effective variable dose (Y) is $1 - R^2 = 0.952$. This means that 9.3% of the chromosome aberration frequency dose (Z) can be predicted by the effective dose (Y) and 90.7% is affected by other variables outside the model.

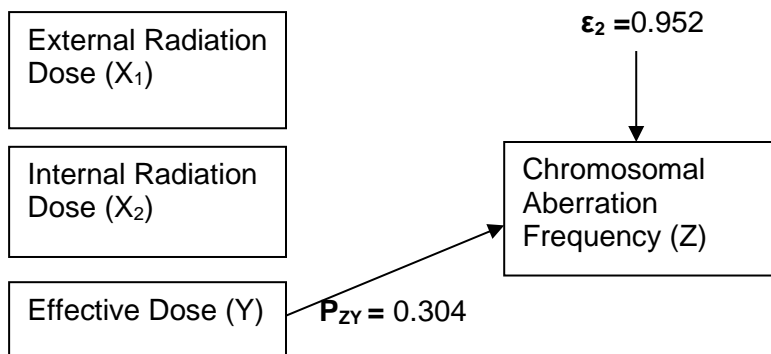


Figure 3. Path Diagram of Structure 2 Path Analysis

Table 3. Overall Simultaneous Hypothesis Test Results of Structure 2

Model	R	R ²	$\epsilon_2 = \sqrt{1 - R^2}$	F	Standard Error	p
Model 1	0.328	0.108	0.944	2.295	0.00308	0.87
Model 2	0.326	0.106	0.945	3.446	0.00305	0.039
Model 3	0.304	0.093	0.952	6.025	0.00305	0.017

Based on the hypothesis test results, the empirical causal model between the external dose (X1), internal dose (X2), and the effective dose (Y) with the frequency of chromosomal aberrations

(Z) can be shown in **Table 4** and **Figure 4**. The structural equations based on the results of hypothesis test are:
 $Y = 0.434X_1 + 0.837X_2 + 0.315$
 $Z = 0.304Y + 0.952$

Table 4. Summary of Hypothesis Test Results After Trimming

Direct Effects Between Variables	Path Coefficient	Standard Error	F-count	P	Conclusion
X ₁ towards Y	0.434	0.013	10.485	0.000/2 = 0.000	Significant
X ₂ towards Y	0.837	0.005	20.209	0.000/2 = 0.000	Significant
Y towards Z	0.304	0.001	2.455	0.017/2 = 0.0085	Significant

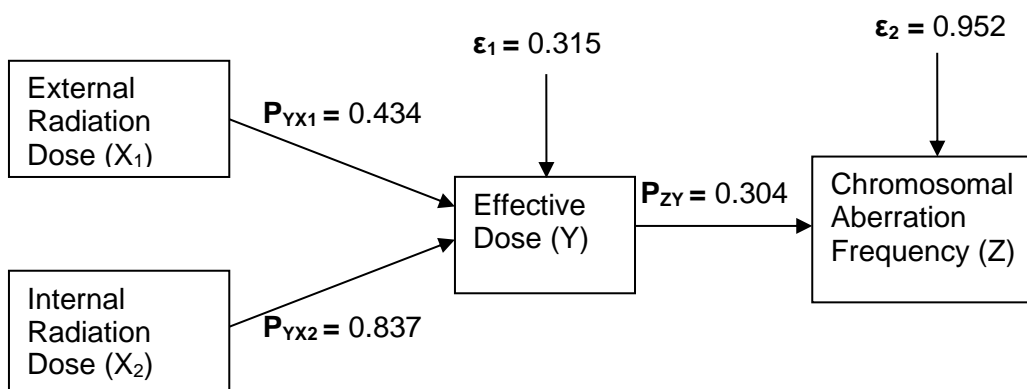


Figure 4. Empirical Causal Model between X₁, X₂, Y, and Z

Model Fit Test

The model fit test is needed to determine whether the proposed hypothetical model is fit or consistent with

the empirical data. The model fit test is done by comparing the theoretical correlation matrix with the empirical correlation matrix. If the two matrices are

identical or appropriate, then the proposed hypothetical model is concluded to be perfectly acceptable. The coefficient of

determination for each structure obtained is presented in **Table 5**.

Table 5. Coefficient of Determination

	Before Trimming	After Trimming
$R_{Y.X_1X_2}^2$ (Structure 1)	0.901	0.901
$R_{Z.X_1X_2Y}^2$ (Structure 2)	0.108	0.093

Determining Chi-Square Statistics

Based on the Chi-Square table with $d = 2$ at the significance level $\alpha = 0.05$ resulted in $X^2_{tab} = X^2_{(0.05;2)} = 5.99$, hence, $X_2 = 0.983903 < X^2_{tab} = 5.99$, therefore H_0 is accepted. It means that the empirical model obtained is appropriate or suitable (model fit) with the data.

The external and internal radiation dose simultaneously and significantly affects the effective dose, amounting to $R^2 = 0.901$ and determination error $\epsilon_1 = 0.315$. This means that 90.1% of the effective dose can be predicted by external and internal radiation dose, 9.9% is affected by other variables besides the model. The external and internal dose of radiation has a significant effect on the chromosome aberrations frequency only through an effective dose of $R^2 = 0.093$ and determination error $\epsilon_1 = 0.952$. It means that 9.3% of the chromosome aberrations frequency can be predicted by external, internal, and effective radiation doses, but 90.7% is affected by other variables outside the model.

The results from this study was: (1) The external dose variable (X_1) did not

have a direct effect on the frequency of chromosome aberrations (Z) but had an indirect determination through the effective dose (Y) of 0.304. Thus, the frequency of chromosomal aberrations in lymphocyte cells, could not be explained by the level of external radiation dose, without going through the effective dose, (2) The internal dose variable (X_2) did not have a direct effect on the frequency of chromosome aberrations (Z) but had an indirect determination through the effective dose (Y) of 0.254. Thus, the frequency of chromosomal aberrations in lymphocyte cells, could not be explained by the internal radiation dose level, without going through the effective dose, (3) The effective dose variable (Y) determined the frequency of chromosomal aberrations in lymphocyte cells (Z). Thus, the frequency of chromosomal aberrations can be explained by the dose, directly equal to 0.304, and (4) The external dose (X_1), internal (X_2), and effective dose (Y) variables had a simultaneous effect on the frequency of chromosome aberrations (Z), of 0.093 or 9.3%. The summary can be seen in **Table 6**.

Table 6. Direct, Indirect, and Total Effect on Endogenous Variables

Exogenous Variables	Path Coefficient	Correlation with endogenous variable (Z)			R-Simultaneous
		direct	indirect (Y)	Total	
External radiation dose (X_1)	0.434	-	0.132	0.132	
Internal radiation dose (X_2)	0.837	-	0.254	0.254	
Effective Dose (Y)	0.304	0.304	-	0.304	
ϵ_1	0.315				
ϵ_2	0.952				
$X_1, X_2 \rightarrow Y$					0.901
X_1, X_2, Y					0.093

Discussion

Natural background radiation sources in question consist of external exposure from cosmic rays (0.4 mSv/year) and terrestrial gamma radiation (0.5 mSv/year), as well as internal exposure, namely inhalation (1.3 mSv/year, especially Radon) and ingestion (0.3 mSv/year)¹.

Exposure to natural background radiation generally originates from soil or rocks and mineral materials containing the thorium series (Th-232), uranium-series (U-238), and potassium-40.³ Areas with high natural background radiation generally contain these substances geographically and are scattered in the environment, thus the radiation levels in the environment are difficult to control. Efforts to reduce external radiation exposure to the body are called, "remedial action".

Efforts to improve the public health degree are necessary by carrying out promotive and preventive activities, by increasing the knowledge of health workers and the inhabitants regarding high natural background radiation. Efforts can be made by training local health workers regarding high natural background radiation and controlling health impacts. After the workers are trained, counseling regarding high natural background radiation, and controlling health impacts can be carried out to the inhabitants in areas with high natural background radiations.

Both external and internal dose levels of natural background radiation will produce an effective dose of radiation that is accepted by the inhabitants. For having information on the exposure to natural background radiation correlation pattern, it is necessary to know the effective dose of radiation received by lymphocyte cells, because the lymphocyte cells can typically change when exposed to high natural ionizing radiation.

Average Effective Dose of Lymphocyte Cells in Botteng Village Inhabitants and Dose Limit Value for Community Members according to BAPETEN Regulation No. 4 of 2013

An area with several chronic exposures that causes the effective dose to

be above the specified level on the inhabitants, is commonly referred to as High Natural Background Radiation (HNBR) (1 mSv/year or 0.12 mSv/year in lymphocyte cell).² Based on the results of univariate analysis of internal, external, and effective doses received by the inhabitants in Botteng Village, it shows that Botteng Village is an area with high natural background radiation.

One of the biological effects caused by ionizing radiation exposure to the body is chromosome aberrations. The chromosome aberrations that can only occur by ionizing radiation exposure are chromosome aberrations in lymphocyte cells. This biological effect can affect the health condition of the inhabitants in the future.

Correlation of Internal and External Radiation Dose towards Effective Dose

Based on the Path analysis in structure 1, the external and internal radiation dose simultaneously and significantly affected the effective dose, amounting to $R^2 = 0.901$ and determination error $\varepsilon_1 = 0.315$. This means that 90.1% of the effective dose can be predicted by external and internal radiation dose, while the other 9.9% was affected by other variables besides the model.

Radioactive materials in the environment expose the human body in two ways, namely exposure to external radiation, and internal radiation exposure. External radiation exposure is ionizing radiation exposure from outside the body. The Path Coefficients $P_{YX1} = 0.434$, every 1 dose of external radiation received contributes 0.434 effective dose of the inhabitant HNBR ($p=0.000$)

The internal radiation exposure is ionizing radiation exposure in which radionuclides contaminated the body through the respiratory tract (inhalation), ingestion (ingestion). The two exposures have different radiation patterns to the body. A path coefficient of $P_{YX2} = 0.837$, means that every 1 dose of internal radiation received contributes to 0.434 effective dose of the inhabitant HNBR ($p=0.000$). The internal dose comes from

radiation exposure sources that enter the body. This resulted in the contribution of the internal dose towards the lymphocyte cells' effective dose, greater than the external dose. The radioactive materials exposure pathways to the body can be through inhalation and ingestion. Factors that can affect the inhalation and ingestion exposure includes the food materials, food sanitation, concentrations of radioactive materials contained in food, consumption patterns, concentrations of radioactive materials in air particulates, home ventilation systems, and respiratory flow rate.⁶⁻⁹ On the other hand, exposure to external radiation is affected by the distance between the radiation source and the body. This resulted in the smaller contribution of the external dose towards the lymphocyte cells' effective dose compared to the internal dose.¹⁰

Correlation of External Radiation Dose to Chromosome Aberrations Frequency in Lymphocyte Cells

The occurrence of chromosomal abnormalities depends on the dose, energy, and type of radiation received.¹¹ The correlation between the probability of a stochastic effect and the dose received can be predicted using the Linear Non-Threshold Hypothesis. The analysis showed that there is no linearity relationship between the external radiation dose and the chromosome aberrations frequency in lymphocyte cells ($p=0.176$). This indicates a phenomenon of radio adaptive response, in peripheral blood lymphocyte cells of people living in areas with high natural background radiation.

Ionizing radiation exposure that occurs in an area with high natural background radiation, Botteng Village, is considered as a low dose radiation exposure. The exposure occurs continuously (chronic). Exposure to low doses of radiation causes non-lethal (non-mutagenic) cell damage and is expressed in subsequent cells. It also make the cell become unstable. The characteristic of unstable cells in the presence of several delayed damages, one of them is the chromosome aberrations. The magnitude of the response and the characteristics of

unstable cells depend on the linear energy transfer and radiation dose, as well as cell type and genetics. External radiation from gamma rays only causes an increase in the aberrations frequency in the 20 – 30th generation of the population after receiving exposure and then decreases towards the background.^{12,13}

Communities in the areas with high natural background radiation receive chronic exposure to low doses of radiation. Human cells can recognize and respond to a very low dose of radiation exposure with several changes, including gene expression. After exposure to a low dose of radiation (adaptive dose), in a short time the cells also exposed to a subsequent radiation in a larger dose (challenge dose). The chromosome aberrations frequency in these cells was lower than in cells that had not received previous low-dose radiation exposure. This shows the existence of cell resistance to exposure to higher doses of radiation in cells that have previously received low doses of exposure.^{12,13}

The existence of a radio adaptive response in lymphocyte cells is proved by a study conducted on residents living in areas with high natural background radiation, in Ramsar, Iran. The study was conducted on peripheral blood lymphocyte cell cultures, which were taken from 15 healthy residents of the area with high natural background radiation as the case group, and 30 healthy residents of the area with normal natural background radiation as the control group. In lymphocyte cell culture, both groups were exposed to external radiation (gamma rays) of 1.5 Gy. It showed that there was a radio adaptive response in residents' lymphocyte cells in the areas of high natural background radiation.¹⁴ This is in line with the results of this study which showed that there was no relationship between external dose and the chromosome aberrations frequency in the inhabitants in Botteng Village.

Correlation of Internal Radiation Dose to Chromosome Aberrations Frequency in Lymphocyte Cells

The analysis results showed that there was no linearity correlation between the internal radiation dose and the

chromosome aberrations frequency ($p=0.376$). Low flux exposure of alpha particles by the board beam does not cross every cell in the radiation environment. In the human body, exposure to a low flux of alpha particles only occurs when the radionuclides contaminated the body (internal radiation). Low flux alpha particles can increase the number of mutations. In addition, it led to the accumulation of the tumor suppressor gene protein p53 (TP53) in a higher percentage of the exposed population compared to the nucleus of cells that received a low flux alpha particle 1 pathway. Thus, it is known that intracellular communication is involved in transmitting several damage signals from cells that receive exposure to cells that do not. Hence this effect can potentially increase the number of damaged cells more than those that were directly exposed to radiation. This means that the internal dose in the human body will affect the chromosome aberrations frequency.⁶

Some damage as a consequence of the response that occurs in several cells is not traversed by an alpha particle directly. Several studies have shown an increase in sister chromatid exchange (SCE) of cells exposed to radiation as a bystander effect. The bystander effect is an effect that occurs in cells that do not receive alpha particle exposure but receive a signal from exposed cells as a result of alpha particle exposure.¹⁵

Theoretically, Sister Chromatid Exchange (SCE) is mediated by the release of extracellular factors and its frequency can be reduced by free radical scavengers.^{15,16} Radionuclides that contaminated the body through inhalation and ingestion, are materials that have unstable atoms, certain atoms can become free radical scavengers due to radiation exposure. Thus, in the up following research, it is necessary to conduct an assessment using the IC50 assay method, on food materials, drink and inhaled air. To determine the presence and effectiveness of certain ingredients that are antioxidants, aiming that those materials can be used to increase resistance to radiation. This study is in line with those efforts that will result in the improvement of public health status in

areas with high natural background radiation.

Effect of Internal Radiation Dose, External Radiation Dose, and Effective Dose on Chromosome Aberrations Frequency in Lymphocyte Cells

Path analysis in structure 2 aimed to see the simultaneous effect of external, internal, and effective doses of lymphocytes on the chromosome aberrations frequency. The correlation between the high dose with the probability of stochastic effects' occurrence can be predicted by using the Linear Non-Threshold Hypothesis. The stochastic effect on humans is caused by high natural background radiation exposure, initiated by a cytogenetic response identified from early biologic markers, which is one of the earliest biologic markers of chromosome aberrations exposure-response. Chromosome Aberrations can be observed through peripheral blood lymphocytes, which are the most sensitive cells to ionizing radiation. These peripheral blood lymphocyte cells also circulate throughout the body, so the damage that occurs in the peripheral blood will represent the damage that occurs in the body.^{17,18}

Based on the analysis, the external and internal dose of radiation has a significant effect on the chromosome aberrations frequency only through an effective dose of $R^2 = 0.093$ and determination error $\epsilon^2 = 0.952$. This means that 9.3% of the chromosome aberrations frequency can be predicted by external, internal, and effective radiation doses, while the other 90.7% is affected by other variables outside the model.

Chronic exposure to low-dose radiation received by people living in areas with high natural background radiation can induce a radio adaptive response. This has been described by several experts.

- a. Low-dose radiation can increase the immune response.¹⁹⁻²¹
- b. Low-dose radiation can reduce DNA damage below spontaneous levels and reduce the likelihood of neoplastic transformation by stimulating cellular detoxification and repair mechanisms.²²⁻²⁴

- c. Exposure to low doses of ionizing radiation can boost the immune system and eliminate the detrimental effects of high doses of radiation.^{25,26}
- d. The immune system of residents in areas with high natural background radiation has been adapted to the naturally high levels of radiation by shifting from Type 1 to Type 2 response to increase anti-inflammatory.^{27,28}
- e. The response of cells exposed to sublethal doses from DNA-damaging agents (non-mutagenic doses of chemicals or radiation), known as conditioning treatment (CT), leads to an increase in resistance to subsequent exposure by higher doses of the similar or other agents, known as challenge treatment (CR).²⁹⁻³¹

Although there is a radio adaptive response and cell repair mechanism, the radiation dose contributes 9.3% to the chromosome aberrations frequency. This indicates the effects of health problems that can occur in the future. Thus, it is necessary to improve the medical record system or clinical data registry inhabitant at health facilities located in areas with high natural background radiations. The purpose of this is to analyze health problems experienced by people in areas with high natural background radiations, in a cohort manner. Therefore, it is necessary to increase cross-sectoral collaboration between stakeholders.

Conclusion

On the basis of the results of the conducted research, the conclusions are as follows:

1. The average dose of external radiation is 5.49 mSv/year, the internal dose is 10.340 mSv/year, and the effective dose of lymphocyte cells is 1.918 mSv/year. The examination results of chromosome aberrations have an average value of 0.00082 from approximately 14.695 metaphase cells observed manually with a microscope.
2. There was a linear relationship between the external radiation dose and effective dose. The Path Coefficients $P_{YX1} = 0.434$ ($p=0.000$)
3. There was a linear relationship between the internal radiation dose and effective dose. The Path Coefficients $P_{YX2} = 0.837$ ($p=0.000$)
4. The external and internal radiation dose simultaneously and significantly affects the effective dose, amounting to $R^2 = 0.901$ and determination error $\epsilon_1 = 0.315$.
5. There was no direct correlation between external, internal radiation dose and chromosome aberrations frequency.
6. The external (indirect correlation = 0.132) and internal dose of radiation (indirect correlation = 0.254) has a significant effect on the chromosome aberrations frequency only through an effective dose, with $R^2 = 0.093$ and determination error $\epsilon_2 = 0.952$.

Ethics approval

Ethics approval from Center for Tech of Radiation Safety, Jakarta (Indonesia).

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Author Contribution

EF analyzed and interpreted the path analysis model and was a major contributor in writing the manuscript. EP performed data collection and analyze data from Center for Technology of Radiation Safety, Nuclear Energy Agency of Indonesia. AS has given advice on the results of the research to be recommended to the relevant agencies and improve the

writing of the manuscript to make it perfect. All authors read and approved the final manuscript.

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