-⊗-

UDC 613.22:577.112.386:577. 164.1:614.876(477.41)

https://doi.org/10.32402/ dovkil2022.03.014

ГІПЕРГОМОЦИСТЕІНЕМІЯ У ДІТЕЙ— ІНДИКАТОР ЕКОЛОГІЧНОГО НЕБЛАГОПОЛУЧЧЯ, ПОВ'ЯЗАНОГО З ЧОРНОБИЛЬСЬКОЮ ЗОНОЮ ВІДЧУЖЕННЯ

Бандажевський Ю.І., Дубова Н.Ф.

HYPERHOMOCYSTEINEMIA IN CHILDREN IS AN INDICATOR OF ENVIRONMENTAL PROBLEMS ASSOCIATED WITH THE CHORNOBYL EXCLUSION ZONE

Ī

¹BANDAZHEVSKYI Yu.I., ²DUBOVA N.F.

¹Ecology and Health Coordination and Analytical Centre, Ivankiv, Ukraine ²Shupyk National Healthcare University of Ukraine, Kyiv he physical development of a person depends on his genetic potential and environmental influences.

Relevant, from a scientific and practical point of view, is the determination of markers of violations of the physiological interaction of endogenous and exogenous factors, leading to the development of severe pathological processes. The sooner it is possible to find a manifestation of a future disease, the more effective will be preventive and therapeutic measures.

The Chornobyl Exclusion Zone (ChEZ) is the territory most contaminated with radioactive elements after the accident at the 4-th power unit of the Chornobyl Nuclear Power Plant (ChNPP) in 1986. Soils and forest trees growing on them contain a huge amount of long-lived radionuclides, of which ¹³⁷Cs, ⁹⁰Sr, and ²⁴¹Am are the most common [1]. Forest fires that constantly occur in the ChEZ contribute to the fact that these radionuclides and their decay products spread with air currents to other areas [2].

Some conventionality of the ChEZ should be noted, since, in the soils and trees, in the territories adjacent to it, the indicated radioactive elements are also determined in large quantities [3]. The population that lives in these ter-

ГІПЕРГОМОЦИСТЕЇНЕМІЯ У ДІТЕЙ – ІНДИКАТОР ЕКОЛОГІЧНОГО НЕБЛАГОПОЛУЧЧЯ, ПОВ'ЯЗАНОГО З ЧОРНОБИЛЬСЬКОЮ ЗОНОЮ ВІДЧУЖЕННЯ

¹Бандажевський Ю.І., ²Дубова Н.Ф. ¹Координаційний аналітичний центр «Екологія і здоров'я», Іванків, Україна ²Національний університет охорони здоров'я України ім. П.Л. Шупика, Київ, Україна

Мета дослідження: визначення участі генетичного фактора (поліморфізм МТНГЯ: С677Т), що впливає на активність основного фермента фолатного циклу – метилентетрагідрофолатредуктази та зовнішньосередовищного впливу, що включає радіаційний чинник, у виникненні стану гіпергомоцистеїнемії у дітей, які проживають поблизу Чорнобильської зони відчуження (ЧЗВ).

Методи дослідження: імунохімічний, математико-статистичний.

Результати. У рамках міжнародних проєктів визначено рівні гомоцистеїну (H_{cy}) та алельних варіантів генетичного поліморфізму МТНFR:C677T у 624 дітей віком 13-17 років із Іванківського та

Поліського районів Київської області. Встановлено, що підвищення рівня Н_с, у крові та збільшення кількості випадків гіпергомоцистеїнемії в обстежених групах дітей пов'язані з лісовими пожежами у ЧЗВ у 2015 та 2020 роках. В умовах вираженого впливу радіоактивних агентів та продуктів горіння лісових дерев питома вага випадків гіпергомоцистеїнемії у групах дітей з алеллю Т поліморфізму MTHFR:677 у геномі і без неї не мала статистичних відмінностей. Індикатором екологічного неблагополуччя у ЧЗВ та прилеглих до неї районах є підвищення понад фізіологічний рівень концентрації H_{cy} у крові дітей підліткового віку з відсутністю у геномі алелі T поліморфізму MTHFR:677. У районах, які постраждали від аварії на Чорнобильській атомній електростанції, необхідне регулярне проведення моніторингових досліджень стану здоров'я дітей з використанням тестів на вміст Н_{су} у крові та визначення генів фолатного циклу.

Ключові слова: гомоцистеїн, гіпергомоцистеїнемія, підлітки, Чорнобильська зона відчуження.

© Бандажевський Ю.І., Дубова Н.Ф. СТАТТЯ, 2022.

№ 3 2022 **Environment & Health** 14

ritories is forced to conduct economic activities, being exposed to the danger of constant radiation exposure.

In many cases, no link can be established between this exposure and severe illness resulting in death or disability.

In our opinion, this is due to a very poor study of the real situation in the territory affected by the Chornobyl accident, when internal organs and the brain are simultaneously incorporated by longlived radionuclides [4].

At the same time, even relatively small concentrations of radionuclides in the body can cause pronounced metabolic disorders, as well as structural and functional changes at the cellular and organ levels [5]. The children's organism is especially sensitive to this influence.

In this regard, the problem of preventive medicine in the areas affected by the Chornobyl accident, in our opinion, is the early diagnosis of these conditions, and hence the selection of markers illustrating the initial stages of the pathological process.

Homocysteine (H_{cy}), a sulfur-containing amino acid, a product of methionine metabolism, should be considered as one of these markers.

It was revealed that an increase in the content of this metabolite in the blood is accompanied by serious disorders in the cardiovascular system, leading to impaired circulation of vital organs [6, 7]. Hyperhomocysteinemia has been associated with oncological diseases, pathology of the nervous system, congenital malformations and impaired pregnancy [8-10].

Mostly hyperhomocysteinemia was recorded in adults, especially the elderly. At the same time, the relationship of this condition with the development of a particular pathological process was assessed.

Monitoring studies were



проблеми чорнобиля

carried out in a number of countries to determine the level of $H_{\rm cy}$ in the blood of physiologically healthy children [11, 12]. However, at the same time, no assessment was made of the influence of internal and external factors on the exchange of this agent.

On the territory bordering the ChEZ, the results of monitoring studies of the child (adolescent) contingent with an assessment of the level of H_{cv} in the blood and the state of the genetic system are very valuable, controlling the main enzymes of the folate cycle (FC), in particular, methylenetetrahydrofolate reductase. A significant decrease in the activity of this enzyme is recorded if the genome contains the risk allele T of the MTHFR:C677T polymorphism [13, 14].

The aim of this study was to determine the involvement of a genetic factor (polymorphism MTHFR:C677T), which affects the activity of the main FC enzyme, methylenete-trahydrofolate reductase, and environmental exposure, including a radiation factor, in the occurrence of a state of hyperhomocysteinemia in children living near the ChEZ.

Material and research methods. The paper ana-

lyzes the results of studies conducted in 2015 in Ukraine within the framework of the projects of the European Commission «Health and **Programmes** Ecological around the Chornobyl Exclusion Zone: Development, training and coordination of health-related projects» and the Regional Council of Rhone-Alpes (France), and also, in January 2022, with the financial support of the public organization «Children of Chornobyl» (France).

624 children from Ivankivskyi and Polesskyi districts of the Kyiv region, located near the ChEZ, were subjected to laboratory examination. After the Chornobyl accident in 1986, the territory of these districts was contaminated with radioactive elements [15]. In particular, studies conducted in 2014 within the framework of the European Union project confirmed the high density of contamination the territory of Ivankivskyi district with ¹³⁷Cs and ⁹⁰Sr radionuclides [3].

In the examined children aged 13-17 years old who attended school, in the morning on an empty stomach, blood was taken from the cubital vein.

The 1-st group included children whose blood sam-Table 1

Number of examined children in age groups

Age, years old	Age, Number of children in groups, persons								
years old	1	2	3						
13	16	49	55						
14	54	73	51						
15	61	56	34						
16	47	72	19						
17	22	11	4						
13-17	200	261	163						

pling was carried out on 04/02/2015, the 2-nd group – 12/18/2015, the 3-rd group – 01/19/2022.

In the course of the ongoing mathematical and statistical analysis, age groups of children were identified (table 1).

The blood samples were analyzed in a laboratory certified according to international quality standards and were agreed with the parents. The content of H_{cy} in the blood was assessed and allelic variants of the MTHFR:C677T genetic polymorphism were determined.

The determination of Hcy in the blood was carried out using the immunochemical method with chemiluminescent detection (ECLIA). Analyzer and test system: Architect 1000 (ABBOT Diagnostics (USA).

The level of H_{cy} in the blood of children over 10 μ mol/l was defined as a state of hyperhomocysteinemia.

During the determination of allelic variants of the genetic polymorphism MTHFR:C677T, the following method was used: Real-time PCR. Analyzer and test system DT-96 detecting cycler; «DNA-Technology» (Russia).

Statistical processing of the obtained results was carried out using the IBM SPSS Statistics 22 program (USA). For the analyzed indicators were calculated the arithmetic mean (M), standard error of the mean (±m), confidence interval of the mean value (95% CI), median (Me), interquartile range (IR), minimum and maximum values of the parameters, percentiles. The hypothesis about the type of distributions was tested (Kolmogorov-Smirnov criterion). All the studied parameters did not correspond to the normal distribution law, and therefore, the non-parametric Mann-Whitney U-test was used to compare the values.

The statistical significance of the indicators was assessed by determining the significance level p using a statistical program.

Student's t-test was used to compare relative scores. The critical confidence level of the null hypothesis (p) was taken as 0.05.

Table 2

The number and proportion of cases of hyperhomocysteinemia ($H_{cy} > 10.0 \ \mu mol/l$) in the age groups of children from Ivankivskyi and Poliskyi districts

Λαο	Gro	up 1	Gro	up 2	Group 3		
Age, years old	Absolute number (n)	Percent- age, %	Absolute number (n)	Percent- age, %	Absolute number (n)	Percent- age, %	
13	6	37.50	32	65.31	32	58.18	
14	27	50.00	57	78.08	35	68.63	
15	32	52.46	42	75.00	24	70.59	
16	25	53.19	56	77.78	15	78.95	
17	8	36.36	11	100.0	4	100.0	
13-17	98	49.00	198	75.86	110	67.48	

Results and its discussion. In the 1-st group of children (13-17 years old), the proportion of cases of hyperhomocysteinemia was significantly less than in the 2-nd and 3-rd groups. There were no statistical differences between groups 2 and 3 (tables 2, 3).

In subgroups of children aged 14, 15, 16 and 17 years old from group 2, the proportion of cases of hyperhomocysteinemia was statistically higher than in similar subgroups from 1-st group. The proportion of cases of hyperhomocysteinemia in subgroups of children aged 16 and 17 years old was 3-rd group, also, was more than in similar subgroups that made up the 1-st group. Between similar age subgroups of the 2-nd and 3-rd groups, significant statistical differences in the proportion of cases of hyperhomocysteinemia were not recorded (tables 2, 3).

The distribution frequency of CC and TT genotypes, CT polymorphism MTHFR:677, in the studied groups of children, as well as between groups, within age subgroups, had no statistical differences (table 4).

The proportion of cases of hyperhomocysteinemia in subgroups of children with and without the T allele from the 2-nd group is statistically more than in similar subgroups from the 1-st group.

In the 3-rd group, in the subgroup of children who do not have the T allele in their genome, the proportion of cases of hyperhomocysteinemia is also significantly more than in the similar sub-

Table 3

Group comparison results by Student's t-test for unrelated populations

Age,	Comparison groups								
years old	[1-2]	[1-3]	[2-3]						
13	t = 2.00; p = 0.052900	t = 1.50; p = 0.143152	t = 0.75; p = 0.456350						
14	t = 3.36; p = 0.001175	t = 1.98; p = 0.052311	t = 1.17; p=0.246696						
15	t = 2.61; p = 0.010921	t = 1.80; p = 0.078091	t = 0.45; p=0.651498						
16	t = 2.80; p = 0.006400	t = 2.17; p = 0.036188	t = 0.11; p=0.912073						
17	t = 6.20; p = 0.000013	t = 6.20; p = 0.000013	-						
13-17	t = 6.09; p = 0.000001	t = 3.63; p = 0.000359	t = 1.85; p =0.065104						



HYPERHOMOCYSTEINEMIA IN CHILDREN IS AN INDICATOR OF ENVIRONMENTAL PROBLEMS ASSOCIATED WITH THE CHORNOBYL EXCLUSION ZONE

1Bandazhevskyi Yu.I., 2Dubova N.F.

1Ecology and Health Coordination and Analytical Centre, Ivankiv, Ukraine

2Shupyk National Healthcare University of Ukraine, Kyiv

The aim of study: determination of the genetic factor's involvement (MTHFR:C677T polymorphism), which affects the activity of the main enzyme of the folate cycle, methylenetetrahydrofolate reductase, and environmental exposure, including the radiation factor, in the occurrence of hyperhomocysteinemia in children which living near the Chornobyl Exclusion Zone (ChEZ).

Research methods. Immunochemical, mathematical and statistical.

Results. Within the framework of international projects, the levels of homocysteine (Hcy) and allelic variants of the genetic polymorphism MTHFR:C677T were determined in 624 children aged 13-17 years old from Ivankivskyi and Poliskyi districts of the Kyiv region. It has been established that an increase in the level of Hcy in the blood and

an increase in the number of cases of hyperhomocysteinemia in the examined groups of children are associated with forest fires in the ChEZ in 2015 and 2020. Under conditions of pronounced exposure to radioactive agents and combustion products of forest trees, the proportion of cases of hyperhomocysteinemia in groups of children with and without the Tallele of the MTHFR:677 polymorphism in the genome did not have statistical differences. An indicator of ecological trouble in the ChEZ and adiacent areas is an increase. above the physiological level, in the concentration of Hcy in the blood of adolescent children with the absence of the Tallele of the MTHFR:677 polymorphism in the genome.

Conclusion. In areas affected by the accident at the Chornobyl nuclear power plant, it is necessary to constantly conduct regular monitoring studies of the health of children, using tests for the content of H_{cy} in the blood and the state of the folate cycle genes.

Keywords: homocysteine, hyperhomocysteinemia, folate cycle genes, adolescent children, Chornobyl exclusion zone.

group from the 1-st group (tables 5, 6). Between the 2-nd and 3-rd groups, statistical differences in these parameters were not found.

Only in the 1-st group, the proportion of cases of hyperhomocysteinemia in the subgroup of children with the T allele in the genome is significantly more than in the subgroup without the T allele. No statistical differences were found in the 2-nd and 3-rd groups (tables 5, 6).

The proportion of cases of hyperhomocysteinemia in subgroups of children aged

15 and 17 years old with the T allele from group 2 was statistically greater than in similar subgroups of group 1.

The proportion of cases of hyperhomocysteinemia in the subgroup of children with the T allele at the age of 16 from the 3-rd group was significantly more than in the same subgroup from the 1-st and 2-nd groups (tables 5, 6).

The proportion of cases of hyperhomocysteinemia in the subgroups of children aged 14, 16 and 17 years old, with the C/C MTHFR:677 genotype from the 2-nd group was

statistically more than in similar subgroups of the 1-st group. In the subgroups of children aged 14 and 17 years old from the 3-rd group, who also did not contain the T allele in the genome, the proportion of cases of hyperhomocysteinemia was more than in similar subgroups of the 1-st group (tables 5, 6).

In group 1, in the subgroup of 14-years-old children with the T allele, the proportion of cases of hyperhomocysteinemia was statistically more than in the subgroup of children of the same age with

Table 4

Number and distribution frequency of MTHFR:C677T genotypes in groups and age subgroups of children from Ivankivskyi and Poliskyi districts depending on FC genotypes

Age,	TT+CT MTHFR:677						CC MTHFR:677					
years	Gro	up 1	Gro	up 2	Gro	up 3	Gro	up 1	Gro	up 2	Gro	up 3
old	Abs.	%	Abs.	%	Abs.	%	Abs.	%	Abs.	%	Abs.	%
13	6	37.50	28	57.14	31	56.36	10	62.50	21	42.86	24	43.64
14	26	48.15	39	53.42	28	54.90	28	51.85	34	46.58	23	45.10
15	27	44. 26	25	44.64	20	58.82	34	55.74	31	55.36	14	41.18
16	28	59.57	40	55.56	7	36.84	19	40.43	32	44.44	12	63.16
17	11	50.00	5	45.45	0	0.0	11	50.00	6	54.55	4	100.0
13-17	98	49.00	137	52.49	86	52.76	102	51.00	124	47.51	77	47.24

the absence of this allele in the genome.

A similar situation was revealed in the 3-rd group in the subgroup of children of 16 years of age (tables 5, 6).

In group 2, there were no statistical differences between subgroups of children of the same age, with and without the T allele in the genome.

The level of H_{cy} in the blood of children from the 1-st group was significantly lower than in the blood of children from the 2-nd and 3-rd groups (tables 7, 8). At the

Table 5 steinemia (H_{cv}>10 µmol/l)

The number and proportion of cases of hyperhomocysteinemia ($H_{cy}>10~\mu mol/l$) in groups and age subgroups of children from Ivankivskyi and Poliskyi districts, depending on the genotypes of FC

Age,	TT+CT MTHFR:677						CC MTHFR:677					
years	Gro	up 1	Gro	up 2	Gro	up 3	Gro	up 1	Gro	up 2	Gro	up 3
old	Abs.	%	Abs.	%	Abs.	%	Abs.	%	Abs.	%	Abs.	%
13	3	50.00	19	67.86	21	67.74	3	30.00	13	61.90	11	45.83
14	18	69.23	31	79.49	20	71.43	9	32.14	26	76.47	15	65.22
15	15	55.56	21	84.00	13	65.00	17	50.00	21	67.74	11	78.57
16	18	64.29	34	85.00	7	100.0	7	36.84	22	68.75	8	66.67
17	4	36.36	5	100.0	-	-	4	36.36	6	100.0	4	100.0
13-17	58	59.18	110	80.29	61	70.93	40	39.22	88	70.97	49	63.64

Table 6
The results of statistically significant differences when comparing the proportion of hyperhomocysteinemia in the analyzed groups in children with genetic polymorphisms MTHFR:677 (CT, TT) and MTHFR:677

Comparison groups	Age, years old	Genotypes	Student's t-test	Significance level, p
1-2	13-17	TT+CT TT+CT	3.68	0.000319
1-2	13-17	CC	5.02	0.000002
1-3	13-17	CC	3.34	0.001229
1-1	13-17	CC TT+CT	2.74	0.007368
1-2	15	TT+CT	2.36	0.024295
1-2	17	TT+CT TT+CT	4.39	0.004623
1-3	16	TT+CT TT+CT	3.95	0.000688
2-3	16	TT+CT TT+CT	2.65	0.011526
		TT+CT CC		
1-2	14	CC	3.88	0.000496
1-2	16	CC	2.32	0.028624
1-2	17	CC	4.39	0.003199
1-3	14	CC	2.49	0.021257
1-3	17	CC	4.39	0.007095
1-1	14	CC TT+CT	2.93	0.007266
3-3	16	CC TT+CT	2.45	0.030653

same time, there were no differences between the 2-nd and 3-rd groups.

The level of H_{cy} in the blood of children aged 14, 15, 16 and 17 years old from the 2-nd group was statistically higher than in the blood of children from subgroups of the same age included in the 1-st group.

The level of H_{cy} in the blood of children aged 14 and 15 from the 3-rd group was also higher than in the blood of children of the same age from the 1-st group (tables 7, 8).

Among children aged 15 years old with the T allele in the genome, the level of H_{cy} in the blood was higher in children of the 2-nd group compared to the 1-st group.

Children aged 14, 15 and 16 years old who are not carriers of the T allele, included in 1-st group, had a lower level of H_{cy} in the blood, compared with children of the same age from 2-nd group. At the same time, children aged 14 and 15 years old from the 1-st group had a lower level of H_{cy} in the blood in comparison with children of the same age from the 3-rd group.

In children with the T allele in the genome, at the age of 14 from the 1-st group, as well as in children at the age of 16 from the 2-nd group,

the level of H_{cy} in the blood was higher than in the blood of children with the CC genotype (tables 9-12).

The obtained results indicate that the indicators of the content of H_{cy} in the blood and the proportion of cases of hyperhomocysteinemia in children had different values depending on the time of the study.

The 1-st group of children was examined on 04/02/2015 – before the fires in the ChEZ forest, the 2-nd group of children was examined on 12/18/2015 – after the forest fires in the ChEZ in spring and summer 2015 [1], the 3-rd group of children was examined on January 19, 2022, after forest fires in a large area of the ChEZ in the spring of 2020 [16].

In the analyzed groups and their age subgroups, there were no statistical differences in the frequency of distribution of CC and TT genotypes, CT polymorphism MTHFR:677.

The proportion of cases of hyperhomocysteinemia and the level of $H_{\rm cy}$ in the blood in the 2-nd and 3-rd groups were more pronounced, in comparison with the 1st group. There were no statistical differences between groups 2 and 3.

In most of the individual age subgroups that made up these groups, a similar relationship was found. The most informative in this regard was the subgroup of children aged 14 years old.

The largest number of statistical differences were found in the age subgroups of children with the absence of the T allele in the genome.

It should be noted that if the values of the proportion of cases of hyperhomocysteinemia in the 1-st group in the subgroup of children with the T allele are statistically more significant than in the subgroup of children without the T allele, then in the 2-nd and 3-rd groups there were no statistical differences between these subgroups.

The same applies to most age subgroups with these genotypes.

Thus, in the groups of children from the 2-nd and 3-rd groups, compared with the 1st group, the proportion of cases of hyperhomocysteinemia, to a greater extent, is due to the presence in the genome of the homozygous variant of the neutral allele C of the MTHFR:677 polymorphism. If in the 1-st group the state of hyperhomocysteinemia associated with the C/C MTHFR:677 genotype was registered in 40 cases, or in 20.0% of the number of children in the group, then in the 2-nd group - in 88 cases, or in 33.72% of the number of children in the group, which is significantly more (t=3.37; p=0.001007), in the 3-rd group – in 49 cases, or 30.06% of the number of children in the group, which, also, significantly more than in the 1st group (t=2.20; p=0.030439).

We are inclined to believe that the increase in the level of homocysteine in the blood of children in the Ivankivskyi and Poliskyi districts, who made up the 2-nd and 3-rd groups of the survey, registered during the ongoing research, is a consequence forest fires in the ChEZ in the spring and summer of 2015 year [1], as well as in the summer of 2020 [16].

Table 7
Statistical characteristics of H_{cy} levels (µmol/l)
in examined children from Ivankivskyi and Poliskyi
districts by age

Age, years	G	Group 1	C	Group 2	Group 3		
old	Ме	IR	Me	IR	Ме	IR	
13	9.43	7.96-11.46	10.88	9.31-12.37	10.80	9.10 – 12.00	
14	9.92	8.20-11.94	12.12	10.12-16.24	11.80	9.20 – 15.50	
15	10.18	7.88-12.21	11.96	9.87-13.51	12.10	9.05 – 14.07	
16	10.20	9.03-13.50	11.65	10.40-14.29	12.30	10.10 – 15.00	
17	9.38	8.32-13.36	13.20	12.58-17.44	13.40	11.43 – 15.23	
13-17	9.90	8.24-12.51	11.80	10.12-13.80	11.40	9.20 – 13.70	

Table 8

The results of statistically significant differences when comparing the levels of $H_{\rm cy}$ (µmol/l) in the blood of children in the analyzed groups

Comparison groups	Age, years old	Comparison group size	Average rank	Mann-Whitney U test value, significance level, p
1	13-17	200	188.12	U = 17524.0;
2		261	263.86	p = 0.0001
1	13-17	200	162.59	U = 12418.0;
3		163	205.82	p = 0.0001
1	14	54	49.63	U = 1195.0;
2		73	74.63	p = 0.0001
1	14	54	45.59	U = 977.0;
3		51	60.84	p = 0.010
1	15	61	49.52	U = 1129.5;
2		56	69.33	p = 0.002
1	15	61	42.72	U = 715.0;
3		34	57.47	p = 0.012
1	16	47	50.26	U = 1234.0;
2		72	66.36	p = 0.013
1	17	22	13.93	U = 53.5;
2		11	23.14	p = 0.010

Radioactive elements and combustion products of trees, with air currents, spread from the fire sites to the territories adjacent to the ChEZ. As a result, the organism of the children living there was subjected to longterm radiation-toxic effects, which resulted in a violation of the functioning of the enzyme systems responsible for the exchange of sulfurcontaining amino acids methionine and H_{cy} . Therefore, despite the fact that the genetic factor, in the form of the T allele of the MTHFR:677 polymorphism, played a significant role in increasing the number of cases of hyperhomocysteinemia, directly affecting the activity methylenetetrahydrofolate reductase, an increase in the level of H_{cy} in the blood occurred even in children without this mutation.

Statistical characteristics of $H_{\rm cy}$ levels (µmol/l) in the examined children of the 1-st group, taking into account the genetic polymorphisms of FC

Age,	MTHF	R:677 (CT + TT)	MTHFR:677 (CC)			
years	Number	H _{cy} , μmol/l		Number	H _c	_y , μmol/l	
old	of cases	Me	IR	of cases	Ме	IR	
13	6	10.22	10.22 8.19-12.00		9.18	7.76-11.50	
14	26	11.41	8.59-15.14	28	9.34	7.30-10.56	
15	27	10.18	7.96-14.01	34	9.94	7.57-11.37	
16	28	10.84	9.10-14.12	19	9.39	8.49-10.89	
17	11	9.39	7.37-22.17	11	9.37	8.94-13.23	
13-17	98	10.38	8.36-14.02	102	9.38	8.02-11.13	

Table 10 Statistical characteristics of H_{cy} levels ($\mu mol/l$) in the examined children of the 2-nd group, taking into account the genetic polymorphisms of FC

Age,	MTH	FR:677	(CT + TT)	MTHFR:677 (CC)			
years	Number		_{cy} , μmol/l	Number	H _c	_y , μmol/l	
old	of cases	Me	IR	of cases	Me	IR	
13	27	11.21	11.21 9.42-12.31		10.86	8.98-12.46	
14	39	11.84	10.10-17.00	34	12.13	10.00-14.40	
15	25	12.10	10.84-14.52	31	11.45	9.25-13.28	
16	40	12.60	10.90-17.71	32	11.06	9.78-12.26	
17	5	13.23 13.00-27.44		6	12.83	12.01-15.97	
13-17	136	12.04	10.53-15.0	125	11.41	9.56-13.08	

Table 11 Statistical characteristics of $\mathbf{H}_{\underline{c}y}$ levels ($\mu mol/l$) in the examined children of the 3-rd group, taking into account the genetic polymorphisms of FC

Age, years	MTHF	R:677	(CT + TT)	MTHFR:677 (CC)			
	Number	H _c	_y , μmol/l	Number	H _c	_y , μmol/l	
old	of cases	Me	IR	of cases	Me	IR	
13	31	11.0 9.30-11.90		24	9.65	8.85-12.75	
14	28	11.85 9.35-15.53		23	11.10	9.20-15.50	
15	20	12.30	8.88-14.58	14	11.95	10.33-14.70	
16	7	12.50	12.10-16.80	12	10.20	8.43-13.73	
17	0	0.0 0.0		4	13.40	11.43-15.23	
13-17	86	11.60	9.38-13.70	77	11.20	9.05-14.40	

In recent years, the environmental situation around the ChEZ has deteriorated significantly. This is due to the increased frequency of forest fires affecting large areas of the ChEZ, and, beyond its borders, as well as the difficult economic situation, as a result of which the population is forced to use wood containing radioactive elements in economic activities, including for heating homes and cooking food.

Since the child's body is very sensitive to changes in the state of the environment, it is a sensitive indicator of the appearance of pathogenic factors in it.

In this regard, it is necessary to constantly conduct regular monitoring studies of the health status of children. in-depth research methods that allow assessing the state of the vital metabolic systems of the body. Identification of unfavorable external environmental factors, and determination of markers of their pathogenic effects on the body, will allow organizing effective preventive measures and preventing the development of serious diseases.

An increased level of H_{cv} in the blood of most adolescents indicates a violation of the development of their body, as well as serious environmental problems in areas bordering the ChEZ.

As a result of the conducted studies, it is possible to make a reasonable statement that an increase in the level of Hcy in a group of adolescent children from areas bordering the ChEZ, who do not have the risk allele T of the MTHFR:677 genetic polymorphism in their genome, may be an indicator of environmental distress associated with the spread of radioactive elements and wood combustion products.

Cases of carriage of the T allele of the MTHFR:677 polymorphism cannot be used to assess the state of



the environment near the ChEZ, since this genetic mutation, in itself, has a direct negative effect on the activity of methylenetetrahydrofolate reductase.

Conclusions

In groups of children aged 13-17 years old living in the lvankivskyi and Poliskyi districts of the Kyiv region, an increase in the level of H_{cy} in the blood and an increase in the number of cases of hyperhomocysteinemia is associated with forest fires in the ChEZ in 2015 and 2020.

Under conditions of pronounced exposure to radioactive agents and combustion products of forest trees, the proportion of cases of hyperhomocysteinemia in groups of children with and without the T allele of the MTHFR:677 polymorphism in the genome did not have statistical differences.

An indicator of ecological trouble in the ChEZ and adjacent areas is an increase, above the physiological level, in the concentration of H_{cy} in the blood of adolescent children with the absence of the T allele of the MTHFR:677 polymorphism in the genome.

In the areas affected by the Chornobyl accident, it is necessary to constantly conduct regular monitoring studies of the health status of children, using tests for the content of H_{cy} in the blood and the state of FC genes.

ЛІТЕРАТУРА

1. Bandazhevskyi Yu.I., Dubovaya N.F. Forest fires in the Chornobyl exclusion zone and children's health. PI Coordination and Analytical Center «Ecology and health» (Ivankiv, Ukraine). Kyiv: Aliant LLC, 2021. 44 p.

2. Азаров С.І., Сидоренко В.Л., Руденко О.В. Вивчення процесу надходження радіонуклідів у повітря при пожежах у лісах зони відчуження ЧАЕС. Проблемы пожарной безопасности: сб. науч. тр. К., 2011. Вып. 30. С.16-23.

3. Bandazhevskyi Yu.I., Dubovaya N.F. Chornobyl catastrophe and childrens health. 35 years of world tragedy. PI Coordination and Analytical Center «Ecology and health» (Ivankiv, Ukraine). Kyiv: Aliant LLC, 2022. 158 p.

4. Bandazhevskyi Yu.I. Chronic Cs-137 incorporation in children's organs. *Swiss Medical Weekly.* 2003. Vol. 133. P. 488-490.

5. Бандажевский Ю.И., Дубовая Н.Ф., Бандажевская Г.С. и др. Чернобыль 25 лет: инкорпорированные радионуклиды Сs-137 и здоровье людей. Под ред. Ю.И. Бандажевского. Координационный аналитический центр «Экология и здоровье». К., 2011.

6. Ganguly P., Alam S. Role of homocysteine in the development of cardiovascular disease. *Nutr J.* 2015. Vol. 14 (6). P. 2-10. doi: 10.1186/1475-2891-14-6

7. Antoniades C., Antonopoulos A., Tousoulis D., Marinou K., Stefanadis C. Homocysteine and coronary atherosclerosis: from folate fortification to the recent clinical trials. *European Heart J.* 2009. Vol. 30 (1). P. 6-15. doi: 10.1093/eurheartj/ehn515

8. Varshney K.K., Gupta J.K., Mujwar S. Homocysteine in-

Table 12

Results of statistically significant differences when comparing H_{cy} levels (µmol/I) in the blood in the analyzed groups of children with genetic polymorphisms MTHFR:677 (CT+TT) and MTHFR:677 (CC)

Groups compa- risons	Age, years old	Genotypes	Number of cases	rank	Mann-Whitney U test value, significance level, p
1	13-17	(CT, TT, CC)	200	188.12	U = 17524.0;
2	13-17	(CT, TT, CC)	261	263.86	p = 0.0001
1	13-17	(CT, TT, CC)	200	162.59	U = 12418.0;
3	13-17	(CT, TT, CC)	163	205.82	p = 0.0001
1	13-17	CC	102	88.48	U = 3771.5;
1	13-17	CT+TT	98	113.02	p = 0.003
2	13-17	CC	125	118.37	U= 6921.5;
2	13-17	CT+TT	136	142.61	p = 0.010
1	13-17	CC	102	89.13	U = 3838,0;
2	13-17	CC	125	134.30	p = 0.0001
1	13-17	CC	102	76.87	U = 2588.0;
3	13-17	CC	77	107.39	p = 0.0001
1	13-17	CT+TT	98	100.80	U = 5027.5;
2	10 17	CT+TT	136	129.53	p = 0.001
1	15	CT+TT	27	21.93	U = 214.0;
2	15	CT+TT	25	31.44	p = 0.024
1	14	CC	28	21.38	U = 192.5;
2	17	CC	34	39.84	p = 0.0001
1	14	CC	28	20.32	U =163.0;
3	17	CC	23	31.00	p = 0.003
1	15	CC	34	27.96	U = 355.5;
2	15	CC	31	38.53	p = 0.024
1	15	CC	34	21.21	U = 126.0;
3		CC	14	32.50	p = 0.011
1	6	CC	19	20.00	U = 190.0;
2		CC	32	29.56	p = 0.026
1	14	CC	28	21.52	U = 192.5;
1	14	CT+TT	26	33.94	p = 0.0001
2	16	CC	32	29.56	U = 418.0;
2	10	CT+TT	40	42.05	p = 0.012

duced neurological dysfunctions: A link to neurodegenerative disorders. *Int J Med Res Health Sci.* 2019. Vol. 8 (4). P. 135-146.

9. Keshteli A., Baracos V., Madsen K. Hyperhomocysteinemia as a potential contributor of colorectal cancer development in inflammatory bowel diseases: A review. *World J Gastroenterol.* 2015. Vol. 21 (4). P. 1081-1090. doi: 10.3748/wjg.v21.i4.1081

10. Botto L.D., Yang Q. 5,10-Methylenetetrahydro-folate reductase gene variants and congenital anomalies: a HuGE review. *Am. J. Epidemiol.* 2000. Vol. 151 (9). P. 862-877. doi: 10.1093/oxfordjournals.aje. a010290.

11. Caldeira-Araъjo H., Ramos R., Florindo C., Rivera I., Castro R., Tavares de Almeida I. Homocysteine Metabolism in Children and Adolescents: Influence of Age on Plasma Biomarkers and Correspondent Genotype Interactions. Nutrients. 2019. Vol. 11, № 646. https://doi.org/10.3390/nu1 1030646

12. Must A., Jacques P.F., Rogers G., Rosenberg IA., Selhub J. Serum Total Homocysteine Concentrations in Children and Adolescents: Results from the Third National Health and Nutrition Examination Survey (NHANES III). The Journal of Nutrition. 2003. Vol. 133, Issue 8. P. 2643-2649.

13. Љkovierovб H., Vidomanovб E., Mahmood S. et al. The molecular and cellular effect of homocysteine metabolism imbalance on human health. *Int J Mol Sci*. 2016. Vol. 17 (10). P. 1710-1733. doi:

10.3390/ijms17101733. 14. Gariglio L., Riviere S., Morales A., Rafael P. Comparison of homocysteinemia and MTHFR677CT polymorphism with Framingham Coronary Heart Risk Score. *Archs Cardiol Mex.* 2014. Vol. 84 (2). P. 71-78. doi: 10.1016/j.acmx.2013.12.00 6. Epub 2014 May 1.

15. Загальнодозиметрична паспортизація та результати ЛВЛ-моніторингу у населених пунктах України, які зазнали радіоактивного забруднення після Чорнобильської катастрофи. Дані за 2011 рік. Збірка 14. МОЗ України, НАМН України, МНС України, ДАЗВ, ДУ «ННЦРМ НАМН України», НД ІРЗ АТН України. К., 2012. 99 с.

ko P.A., Vasylenko V.V. et al. Results of WBC – monitoring of firefighters participating in response to Chornobyl forest fires in April-May 2020. *Problems of Radiation Medicine and Radiobiology*. 2020. Vol. 25. P. 177-187. doi: 10.33145/23048336202055

16. Bazyka D.A., Fedir-

P. 177-187. doi: 10.33145/23048336202025 177187.

REFERENCES

1. Bandazhevskyi Yu.I. and Dubova N.F. Forest Fires in the Chornobyl Exclusion Zone and Children's Health. PI Coordination and Analytical Center «Ecology and Health» (Ivankiv, Ukraine). Kyiv: Aliant LLC; 2021: 44 p.

Kyiv: Aliant LLC; 2021: 44 p. 2. Azarov S.I., Sydorenko V.L. and Rudenko O.V. Vyvchennia protsesu nadkhodzhennia radionuklidiv u povitria pry pozhezhakh u lisakh zony vidchuzhennia ChAES [Study of the Process of Radionuclides' Release into Atmosphere During Fires in the Forests of the ChAES]. In: Problemy pozharnoy bezopasnosti: sb. nauch. tr. [Problems of Fire Safety: Collection of scientific works]. Kyiv; 2011; 30 : 16-23 (in Ukrainian).

3. Bandazhevskyi Yu.I. and Dubova N.F. Chornobyl Catastrophe and Childrens Health. 35 Years of World Tragedy. PI Coordination and Analytical Center «Ecology and Health» (Ivankiv, Ukraine). Kyiv: Aliant LLC; 2022: 158 p.

4. Bandazhevskyi Yu.I. Chronic Cs-137 Incorpora-

tion in Children's Organs. Swiss Medical Weekly. 2003; 133: 488-490.

5. Bandazhevskyi Yu.I., Dubovaya N.F., Bandazhevskaya G.S. et al. Chernobyl 25 let: inkorporirovannye radionuklidy Cs-137 i zdorove lyudei. Koordinatsionnyi analiticheskiy tsentr «Ekologiya i zdorove» [Chernobyl 25 Years: Incorporated Cs-137 Radionuclides and Human Health. Coordinating Center «Ecology and Health»]. Kiev; 2011: 156 p. (in Russian).

6. Ganguly P. and Alam S. Role of Homocysteine in the Development of Cardiovascular Disease. *Nutr J.* 2015; 14 (6): 2-10. doi: 10.1186/1475-2891-14-6.

7. Antoniades C., Antonopoulos A., Tousoulis D., Marinou K. and Stefanadis C. Homocysteine and Coronary Atherosclerosis: from Folate Fortification to the Recent Clinical Trials. *European Heart J.* 2009; 30 (1): 6-15. doi: 10.1093/eurheartj/ehn515

8. Varshney K.K., Gupta J.K. and Mujwar S. Homocysteine Induced Neurological Dysfunctions: A Link to Neurodegenerative Disorders. Int J Med Res Health Sci. 2019; 8 (4): 135-146.

9. Keshteli A., Baracos V. and Madsen K. Hyperhomocysteinemia as a Potential Contributor of Colorectal Cancer Development in Inflammatory Bowel Diseases: A Review. World J Gastroenterol. 2015; 21(4): 1081-1090. doi: 10.3748/wjg.v21.i4.1081

10. Botto L.D. and Yang Q. 5,10-Methylenetetrahydro-folate Reductase Gene Variants and Congenital Anomalies: a HuGE Review. *Am. J. Epidemiol.* 2000; 151 (9): 862-877. doi: 10.1093/oxfordjournals.aje. a010290.

11. Caldeira-Araъjo H., Ramos R., Florindo C., Rivera I., Castro R. and Tavares de Almeida I. Homocysteine Metabolism in Chil-



dren and Adolescents: Influence of Age on Plasma Biomarkers and Correspondent Genotype Interactions. *Nutrients*. 2019; 11:646. https://doi.org/10.3390/nu11030646

12. Must A., Jacques P.F., Rogers G., Rosenberg I.H. and Selhub J. Serum Total Homocysteine Concentrations in Children and Adolescents: Results from the Third National Health and Nutrition Examination Survey (NHANES III), *The Journal of Nutrition*. 2003; 133 (8): 2643-2649.

13. Љkovierovб H., Vidomanovб E., Mahmood S. et al. The Molecular and Cellular Effect of Homocysteine Metabolism Imbalance on Human Health. *Int J Mol Sci.* 2016; 17 (10): 1710-1733. doi: 10.3390/ijms17101733.

14. Gariglio L., Riviere S., Morales A. and Rafael P. Comparison of Homocysteinemia and MTHFR677CT Polymorphism with Framingham Coronary Heart Risk Score. *Archs Cardiol Mex.* 2014; 84 (2): 71-78. doi: 10.1016/j.acmx.2013.12.00 6. Epub 2014 May 1.

Zahalnodozymetrychna pasportyzatsiia ta rezultaty LVL monitorynhu v naselenykh punktakh Ukrainy, yaki zaznaly radioaktyvnoho zabrudnennia pislia Chornobylskoi katastrofy. Dani za 2011 rik. [General Dosimetric Certification and Results of WBC-Monitoring in the Settlements of Ukraine Suffered from the Radioactive Contamination after Chornobyl Catastrophe]. Kyiv; 2012; Iss. 14:99 (in Ukrainian).

16. Bazyka D.A., Fedir-ko P.A., Vasylenko V.V. et al. Results of WBC – monitoring of firefighters participating in response to Chornobyl forest fires in April-May 2020. *Problems of Radiation Medicine and Radiobiology.* 2020; 25: 177-187. doi: 10.33145/23048336202025 177187.

Надійшло до редакції 18.04.2022

УДК 314.44 : 616-005 : 614.876 https://doi.org/10.32402/ dovkil2022.03.023

THE STATE OF DISABILITY IN THE ADULT POPULATION EVACUATED FROM THE 30-KM ZONE OF THE CHORNOBYL NUCLEAR POWER PLANT DUE TO DISEASES OF THE CIRCULATORY SYSTEM Kapustynska O.A.

СТАН ІНВАЛІДНОСТІ ВНАСЛІДОК ХВОРОБ СИСТЕМИ КРОВООБІГУ У ДОРОСЛОГО НАСЕЛЕННЯ, ЕВАКУЙОВАНОГО 13 30-км ЗОНИ ЧОРНОБИЛЬСЬКОЇ АЕС

X

КАПУСТИНСЬКА О.А.

ДУ «Національний науковий центр радіаційної медицини НАМН України», Київ вороби системи кровообігу (ХСК) призводять до високого рівня захворюваності, інвалідності, смертності та значною мірою впливають на збільшення витрат на охорону здоров'я, особливо з постарінням населення [1, 2]. Ключове місце в оцінці здоров'я належить інвалід-

СТАН ІНВАЛІДНОСТІ ВНАСЛІДОК ХВОРОБ СИСТЕМИ КРОВООБІГУ У ДОРОСЛОГО НАСЕЛЕННЯ, ЕВАКУЙОВАНОГО ІЗ 30-км ЗОНИ ЧОРНОБИЛЬСЬКОЇ АЕС Капустинська О.А.

ДУ «Національний науковий центр радіаційної медицини НАМН України», Київ

Ключове місце в оцінці здоров'я населення належить інвалідності з урахуванням її високої поширеності й різноманіття соціально значущих наслідків. Наукові дослідження в Україні і в усьому світі засвідчують глобальну поширеність хвороб системи кровообігу (ХСК) серед населення різних вікових груп. До найбільш несприятливих наслідків Чорнобильської катастрофи для здоров'я та соціального благополуччя належить інвалідизація постраждалого населення.

Мета дослідження: проаналізувати динаміку показників інвалідності дорослого населення, евакуйованого після аварії на ЧАЕС, внаслідок провідних ХСК з акцентом на особливості їх розподілу за групами первинно визнаних інвалідами.

Матеріали та методи дослідження. Проведено ретроспективний аналіз стану первинного виходу на інвалідність внаслідок ХСК серед дорослого евакуйованого населення за даними Державного реєстру України осіб, які постраждали внаслідок Чорнобильської катастрофи. Аналіз інвалідності проводили за даними осіб, первинно визнаних інвалідами, серед дорослого евакуйованого населення залежно від часу та віку на дату аварії, за класами, формами патології та групами. Період спостереження — 1988-2016 роки. Обробка отриманих даних потребувала формування комп'ютерної бази даних на основі Microsoft Excel.

Результати. Згідно з проведеним аналізом ХСК продовжують, як і у попередні післяаврійні роки, посідати перше місце у структурі причин первинної інвалідності у дорослого евакуйованого населення. З часом, що минув після аварії, рівень інвалідності підвищувався порівняно з раннім періодом і досяг найвищих рівнів у пізньому періоді

© Капустинська О.А. СТАТТЯ, 2022.

23 Environment & Health № 3 2022