

Is Exposure to Ionizing Radiation Resulting from the Chernobyl Fallout Associated with Childhood Cardiac Arrhythmia?

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Faire avancer la sûreté nucléaire

Exposure to Ionizing Radiation Resulting from the Chernobyl Fallout and Childhood Cardiac Arrhythmia: A Population Based Study

[Jourdain et al., BMJ Open, 2018, accepted]

ICMPRPR – LONDON – March 15th, 2018

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Context

Nuclear accident of Chernobyl (Ukraine – April 26th, 1986)

Radioactive plume (~ 5,3 EBq) mainly composed of iodine and caesium radioactive



Population radiation exposure

Mainly through ingestion of contaminated products (vegetables, fish products, meat, milk, etc.)

Health effects

Cancerous

Thyroïd cancer especially in children under 5 years old (6 848 cases between 1991 and 2005 in children < 18 years old at the time of the accident): link with ionizing radiation exposure proved [Unscear 2008]

Non cancerous

Many diseases or effects reported (chromosomal abnormalities, reproductive effects, psychological effects, peptic ulcers, autoimmune thyroiditis, lens opacities, cardiovascular effects, etc.): link with ionizing radiation exposure?

[Lazjuk and al., Stem Cells, 1997; Little M.P., Radiat Environ Biophys, 2011]

Context

EXPOSURE TO LOW DOSES OF IONIZING RADIATION

■ Nuclear reactor accident (Chernobyl): of the risk of mortality due to ischemic heart disease and cerebrovascular disease [Ivanov et al., Health Physics, 2006]



• German uranium miners: No association between cardiovascular mortality and exposure to radiation

[Kreuzer, Radiat Environ Biophys, 2006]

• Youri Bandazhevski works: study of non-cancerous diseases presented by children living on the contaminated territories of Belarus according to ¹³⁷Cs whole-body burden

[Bandazhevski Y. and Bandazhevskaya G., 2004]

- → Cardiac rhythm disorders
- → Other pathologies: cataracts, peptic ulcers, anemia, etc.
- → Correlation between the level of ¹³⁷Cs contamination and the occurrence of non-cancerous pathologies according to the authors' conclusions

Context

Studies on low doses ionizing radiation \rightarrow limitations +++ [Kreuzer et al., Mutation Research, 2015]

- Methodology not thorough (equipment used, statistical tests, etc.)
- Uncertain dosimetry estimates
- Adjustement problems on cofactors (tobacco, alcohol, etc.)
- Limited statistical power
- No consideration of potential biases, etc.



Little evidence of an association between exposure to ionizing radiation (<0.5 Gy) and the presence of circulatory diseases



IMPLEMENTATION OF EPIDEMIOLOGICAL STUDY

- ➤ Scientific community → to collect scientific information about the non-cancerous effects induced by chonic exposure to low doses of ionizing radiation
- ➤ **Population** → to answer societal issue relating to health consequences Chernobyl nuclear accident in a sensitive population (children)

Cardiac arrhythmia study

- **Study period**: May 2009 May 2013
- Cross-sectional study with a design exposed unexposed
- **Location** → Bryansk oblast (Russian Federation)
- Objectives
 - What is the prevalence of cardiac arrhythmias in Bryansk oblast (CT vs NCT)?
 - Is caesium 137 associated factor in the occurrence of cardiac arrhythmias?
- **Study population**: 17 697 children aged from 5 to 18 years old



Cardiac arrhythmia study

- Medical examinations for all children
 - 137Cs whole-body activity measurement during 10 min
 - EKG during 10 min
 - Cardiac ultrasound
 - Further medical examinations for some children
 - Holter (21%)
 - Blood test (6%)
 - Enzymatic profile (transaminases, PAL, LDH, γ-GT, CK)
 - ✓ Biochemical profile (Na, K, Ca, P, albumin)
 - ✓ Inflammatory profile (CRP)
 - ✓ Renal profile (urea, creatinine)
 - ✓ Thyroïd profile (TSH, FT4)
 - ✓ Cardiac profile (troponin T, CK-MB)
- **Data collection in schools** by a multidiscicplinay team from the Clinical Diagnostic Centre, Bryansk (Russian Federation)

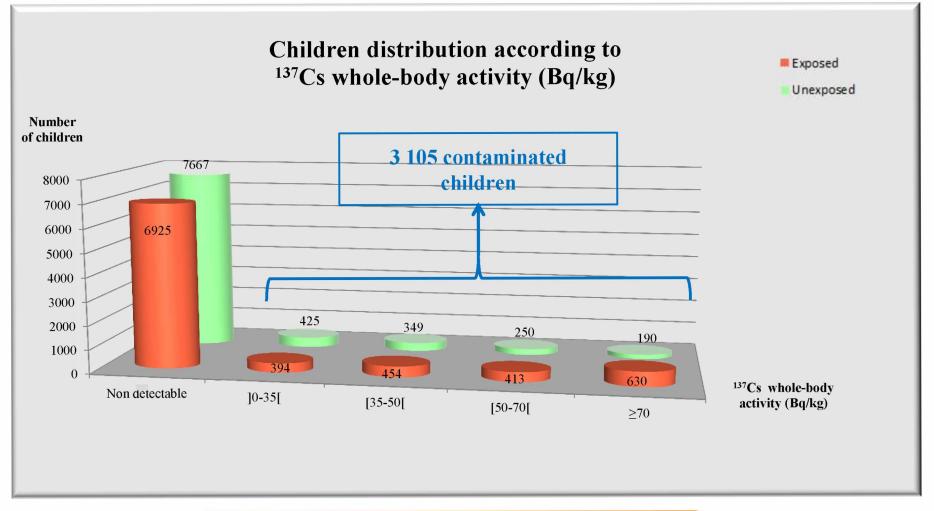
Cardiac arrhythmia study- Population characteristics (1/3)

	CONTAMINATED TERRITORIES (EXPOSED POPULATION)		CONTROLLED TERRITORIES (UNEXPOSED POPULATION)	
CHARACTERISTICS	N = 8 816 (%)		N = 8881	(%)
GENDER				
Male	4 436	(50.3)	4 534	(51.1)
Female	4 380	(49.7)	4 347	(48.9)
AGE				
< 10	3 412	(38.7)	3 057	(34.4)
10 - 13	2 598	(29.5)	3 109	(35.0)
≥ 14	2 806	(31.8)	2 715	(30.6)
DRUG INTAKE				
No	3 933	(44.6)	8 335	(93.9)
Yes	108	(1.2)	257	(2.9)
Do not know	4 775	(54.2)	289	(3.3)

Cardiac arrhythmia study- Population characteristics (2/3)

	CONTAMINATED TERRITORIES (EXPOSED POPULATION)		CONTROLLED TERRITORIES (UNEXPOSED POPULATION)	
CHARACTERISTICS	N = 8816	(%)	N = 8881	(%)
BMI (WHO CLASSIFICATION)				
Underweight	179	(2.0)	204	(2.3)
Normal weight	6 565	(74.5)	6 550	(73.8)
Overweight	1 219	(13.8)	1 218	(13.7)
Obesity	853	(9.7)	909	(10.2)
PLACE OF RESIDENCE				
Rural	2 833	(32.1)	3 165	(35.6)
Urban	5 983	(67.9)	5 716	(64.4)
DIETARY HABITS				
Shop	8 128	(92.2)	7 676	(86.4)
Personal farm (milk, meat, vegetables)	6 448	(73.1)	6 722	(75.7)
Forest (mushrooms, berries, meat)	5 486	(62.2)	7 171	(80.7)
Rivers (fish, crayfish)	3 862	(43.8)	4 237	(47.7)

Cardiac arrhythmia study- Population characteristics (3/3)



COMMITTED EFFECTIVE DOSE

3 081 children < 1 mSv/y 24 children ≥ 1 mSv/y (max = 5,77 mSv/y)



Cardiac arrhythmia study- Diagnosed cardiac pathologies/syndromes

(WHO - ICD – 10^{th} edition) \rightarrow 54 in total

- D15 Benign neoplasm of other and unspecified intrathoracic organs
- I08 Multiple valve diseases except endocarditis
- I34 Nonrheumatic mitral valve disorders
- I35 Nonrheumatic aortic valve disorders
- I36 Nonrheumatic tricuspid valve disorders
- I37 Pulmonary valve disorders
- I42 Cardiomyopathy
- I43 Cardiomyopathy in diseases classified elsewhere
- I44 Atrioventricular and left bundle-branch block
- I45 Other conduction disorders
- I47 Paroxysmal tachycardia
- I49 Other cardiac arrhythmias
- Q20 Congenital malformations of cardiac chambers and connections
- Q21 Congenital malformations of cardiac septa
- Q22 Congenital malformations of pulmonary and tricuspid valves
- Q23 Congenital malformations of aortic and mitral valves
- Q24 Other congenital malformations of heart
- Q25 Congenital malformations of great arteries
- R00 Abnormalities of heart beat

2 526 children with cardiac arrhythmia

Cardiac arrhythmia study - Objective n°1: prevalence estimation

Health status of the study population according to the territory (CT vs NCT)



			TERRITORIES		
			CONTAMINATED	CONTROLLED	
HEALTH Healthy STATUS With cardi	Healthy	7 644	7 527		
	With cardiac arrhythmia	1 172	1 354		

Prevalence estimation over the period 2009 – 2013

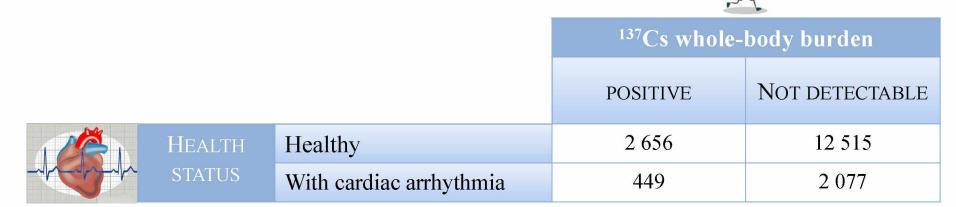
- $P_{\text{exposed}} = 13.3\%$
- $P_{unexposed} = 15.2\%$

Prevalence ratio (PR) = P_{exp}/P_{unexp} = 0.87 [95%IC= 0.84 – 0.90]

Over the period 2009 - 2013, the estimated prevalence of cardiac arrhythmias in CT was significantly lower than in the NCT

Cardiac arrhythmia study - Objective n°1: prevalence estimation

Health status of the study population according to 137Cs whole-body burden



Prevalence estimation over the period 2009 – 2013

- $P_{\text{exposed}} = 14.5\%$
- $P_{unexposed} = 14.2\%$

Prevalence ratio (PR) = P_{exp}/P_{unexp} = 1.02 [95%IC= 0.97 – 1.07]

Over the period 2009 - 2013, the estimated prevalence of cardiac arrhythmia among contaminated children does not differ significantly from the estimated prevalence among children with ¹³⁷Cs contamination not detectable

Cardiac arrhythmia study - Objective n°2: 137Cs= associated factor?

Multivariate analysis (SAS software)

Explanatory variables	OR¥	[_{95%} CI]	P-value
Territory			
Controlled	1		
Contaminated	0.90	[0.81 - 1.00]	0.06

⁴ adjustment for gender, age, drug intake, BMI and ¹³⁷Cs whole-body burden

Explanatory variables	OR¥	[_{95%} CI]	P-value
137Cs whole-body activity (Bq/kg)			
Not detectable	1		
]0 – 35[1.07	[0.89 - 1.29]	
[35 – 50[1.00	[0.81 - 1.23]	
[50 – 70[1.04	[0.82 - 1.31]	
≥ 70	0.96	[0.77 - 1.20]	0.97

[¥] adjustment for gender, age, drug intake, BMI and territory

No evidence of an association between the presence of cardiac arrhythmia and the territory or ¹³⁷Cs whole-body burden

→ 137Cs is not an associated factor in the frame of the study

Cardiac arrhythmia study - Strenghts/ limitations

Strengths

- Study population: 17 697 children
- High participation rate: 91%
- Little missing data (37 for holter and 69 for blood test)
 - → justifications provided
- Conditions gathered to bring out an effet if this one exists



Limitations

- Selection bias
- Memory bias (drug intake)
- Classification bias (no Holter for all children and DL)
- Some factors not considered (blood pressure measurement, alcohol/drug/tobacco consumption, genetic tests, blood test, etc.)

Cardiac arrhythmia study — Conclusions/Perspectives

CONCLUSIONS

- Estimated prevalence over the period 2009-2013
- Statistically significant differences according to the territory (lower prevalence in the contaminated territories)
- No statistically significant differences according to ¹³⁷Cs whole-body burden (contaminated children *vs* children with ¹³⁷Cs contamination not detectable)
- 137Cs is not a factor associated to the presence of cardiac arrhythmia (main hypothesis) in the frame of the study
- Children (with cardiac arrhythmia) care
- Followed up and outpatient care by cardiologists, pediatricians
- 6 children were operated in emergency (2 with the pose of pacemaker)

PERSPECTIVES

- Results communication to Bryansk oblast population
- Lens opacities study (2013 2017)





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