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## ASSESSING OCCUPATIONAL CARCINOGENIC RISKS FOR HEALTH OF WORKERS EMPLOYED AT BLISTER COPPER PRODUCTION ENTERPRISE

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*In order to develop ideological grounds for complex assessment of industrial carcinogenic danger, we performed hygienic assessment of contribution made by working environment factors into carcinogenic risks formation for workers employed at blister copper production. The assessment included predictive values of occupational risks. We determined tumor markers levels for occupations with unacceptable levels of carcinogenic risks. We examined mortality caused by malignant neoplasms. It is shown that non-organic arsenic compounds applied in blister copper production are the main factor causing carcinogenic risks. As we calculated individual carcinogenic risks for 25-year working period we found out that total carcinogenic risks for all basic and supplementary occupations in copper-smelting workshops were within the 4th range (more than  $1.0 \times 10^{-3}$ ). Unacceptable predictive values of carcinogenic risks for 100 % examined occupations appeared already after 5 years of work. We detected excess levels of tumor markers in 73 % of examined workers with occupations characterized with unacceptable values of carcinogenic risks; 19 % of such workers had excess levels of two tumor markers at once. 9 % of examined workers had excess levels of Cyfra 21.1 tumor marker, 14,5 % of workers had high levels of CEA tumor marker, and 59 % of workers had high levels of NSE tumor marker. Intensive mortality indices for male workers employed at copper-smelting workshops as per all localizations amounted to 153.14, whereas they amounted to only 127.25 per 100,000 people for other population groups. Workers employed at blister copper production had higher intensive mortality indices than other people in terms of such nosologies as malignant neoplasms in respiratory organs and chest (86.78 and 47.72 correspondingly), including lung cancer (71.47 and 43.48 correspondingly). The obtained results will help to work out a system for managing carcinogenic risks which will include regulating and controlling activities, managerial activities, technical and technological measures, financial and economic activities, medical and prevention activities plus rehabilitation activities, as well as information and educational activities.*

**Key words:** blister copper production, occupational individual carcinogenic risk, mortality caused by malignant neoplasms, tumor markers, carcinogenic risk management, arsenic, nickel, lead, cadmium.

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Nowadays practically one third of workers in Russia have to work in hazardous and dangerous working environment. As per International Labor Organization assessment, more than 76,000 death cases in Russia are caused by occupational diseases. This situation calls for changes in priorities when planning strategies aimed at population health improvement; namely. the focus needs to be shifted from clinical approach to prevention programs implementation. Therefore, managing occupational risks becomes a most vital task and it means working out a set of activities for preventing adverse impacts exerted by working conditions on health; these activities are to be based on assigning top priority to primary prevention and lowering risk factors influence [16].

P.V. Serebryakov (2007), A.V. Meltser (2008), and some other authors dedicated their research to development of carcinogenic risk assessment methodology. In particular, P.V. Serebryakov was the first to work out an algorithm for calculating individual occupational carcinogenic risks; this algorithm was tested at enterprises which specialized in extracting and processing copper-nickel ores [14]. Issues of occupational exposure to carcinogens, risk assessments and biological monitoring (on the example of arsenic compounds) are also profoundly discussed in foreign scientific literature [17, 18, 20].

Since 2011 Rospotrebnadzor offices and establishments in Sverdlovsk region, together with FBSE "Yekaterinburg medical scientific center for prevention and protection of workers employed at industrial enterprises", have been developing ideological grounds for complex assessment of enterprises' carcinogenic danger [9]. As per results of assessing exposure to carcinogenic factors and working population characteristics, predictive values of individual occupational carcinogenic risk (CR) are calculated; these values are then compared with risks related to non-

production impacts. These CR assessments enable researchers to form a risk group of workers who then undergo additional research on early detection of neoplasms signs during preventive examinations; when it is necessary, such workers have additional examinations in a hospital with further regular medical check-ups. In order to give grounds for production factors contribution into malignant neoplasms (MN) evolvement, research on workers' oncologic mortality is conducted. All the above-stated allows us to justify activities aimed at managing carcinogenic risks.

However, there is a number of methodological problems related to calculation of predictive CR values as its aspects, values and criteria are not confirmed in accordance with the established procedure; there is also an issue of assessing proof of relation between an occupation and cancer evolvement as well as determining tumor formation predictors [7,15].

**Our research goal was** to examine working conditions and assess occupational carcinogenic risks for workers employed at enterprises dealing with blister copper production.

**Data and methods.** We chose a major enterprise in the Urals specializing in blister copper production as our research object; production in copper-smelting workshop at this enterprise combined such technological processes as mix materials drying, smelting in Vanyukov's furnace (PV-1500), and copper converting.

At the first stage of our research we identified the enterprise carcinogenic danger basing on the initial data taken from its sanitary-hygienic certificate. We determined priority carcinogenic factors of production components (raw materials, products, emissions into atmosphere, sewage discharge, air composition in working area etc.) and formed a data bank (on concentrations) for further assessment of occupational carcinogenic health risk. We based our CR calculations on approaches

stated in P 2.1.10.1920-04. "Guidelines on assessment of population health risk under exposure to chemicals which pollute environment" [8] and research conducted by A.V. Meltser and P.V. Serebryakov [5, 14]. CR was calculated for 17 occupations of copper-smelting workshop (drying section, smelting section, and converting section); 420 people were employed at this workshop, namely: drying operators (6 people), carrier operators (15 people), mix materials handlers (2 people), mix materials loaders (39 people), smelters (70 people), casters (30 people), blowers (37 people), crane drivers (31 people), cleaners (52 people), fireproof materials handlers (3 people), foremen (12 people), powder-gas catching devices operators (44 people), pumping units operators (10 people), electro-gas welders (15 people), maintenance fitters (42 people), electricians (25 people), supervisory instruments and automatic devices fitters (5 people), allowing for actual exposure to arsenic, cadmium, lead, beryllium, and benzpyrene (250 working shifts, 8 hours long each).

At the second stage, we determined tumor markers in blood serum of workers who had these occupations; tumor markers included cancer embryonic antigen (CEA) which was a marker of tumors in trachea, bronchial tubes and lungs, gastrointestinal tract, especially large intestine, pancreas, liver, as well as mammary gland, uterine neck and prostate; Cyfra 21.1 which was a marker of lung cancer (mostly epidermoid one, and less frequently adenocarcinoma and other histological types) and urinary bladder; neuron-specific enolase (NSE) which was a marker of lung tumors, leukemia and tumors of neuroectodermal origin [19].

At the third stage we performed retrospective epidemiologic research on mortality caused by malignant neoplasms among workers employed at copper-smelting workshops and having the same occupations for which CR was assessed [6]. Population living in close proximity to the enterprise

was our control group. The research period was 30 years (1976-2005). We calculated intensive mortality indices per 100,000 people of population and workers (distributed as per age and total ones). In addition to the observed mortality in the examined cohorts, we calculated so called expected mortality which was control population mortality standardized as per age. We took age distribution in copper-smelting workshop as a standard. Multiplicity of excess in observed parameters of mortality caused by malignant neoplasms determined the degree of additional risk related to work at the examined production.

**Results and discussion.** Production aerosols are the primary occupational-hygienic hazard factor among all production factors. As ore raw materials are multi-component ones it makes determining dust chemical structure more complicated. Beside main metal, such materials contain a number of carcinogenic substances such as arsenic, nickel, cadmium, hexavalent chromium, and beryllium [1,4,10]. When concentrate is dried and smelted it involves benzpyrene emissions into working area air [2,3]. Dust of mix materials in smelting section contains 0.25-0.6% of arsenic, 0.05-0.09% of lead and cadmium. As for dust in converting section, it contains 0.03-0.09% of arsenic, 0.04-4.38% of lead and less than 0.1% of beryllium.

We detected increased concentrations of lead and arsenic (up to 0.1 and 0.015 mg/m<sup>3</sup>, correspondingly) for some occupations in the course of our research. Cadmium, beryllium and benzpyrene content didn't exceed maximum permissible concentration (MPC). Therefore, working conditions in copper-smelting workshop can be classified as having 2.0-3.1 hazard category.

Individual CR calculation for 25-years long working period revealed that total CR was within the 4th range (more than  $1.0 \times 10^{-3}$ ) for all the examined main and auxiliary occupations of copper-smelting workshop (Table 1). This range is considered

to be unacceptable for occupational groups [8], and it fits in with all the previous research as well as supplements them [1,13].

The greatest CR value was detected at workplaces where dust emissions were the biggest (mix materials handler, mix materials loader, carrier operator, powder-gas catching device operator) and also for some repair occupations (electrician,

maintenance fitter, supervisory instruments and automation devices fitter). The analysis of the obtained results showed that maximum contribution into CR values at all workplaces was caused by exposure to non-organic arsenic compounds (from 84 to 99%). Benzpyrene, beryllium, cadmium and lead content didn't exert any substantial influence on total CR values.

Table 1

Individual carcinogenic risks for workers employed at copper-smelting workshop

Workplace	Carcinogenic risk at 25-year long working period					Total carcinogenic risk
	Carcinogenic substances					
	As	Cd	Pb	Benzpyrene	Be	
Drying operator	$3,2 \times 10^{-3}$	$4,4 \times 10^{-5}$	$4,6 \times 10^{-5}$	–	–	$3,3 \times 10^{-3}$
Carrier operator	$4,2 \times 10^{-3}$	$4,4 \times 10^{-5}$	$7,5 \times 10^{-4}$	–	–	$5,0 \times 10^{-3}$
Mix materials handler	$5,2 \times 10^{-2}$	–	$4,4 \times 10^{-5}$	–	–	$5,3 \times 10^{-2}$
Mix material loader	$7,9 \times 10^{-3}$	$2,2 \times 10^{-5}$	$1,2 \times 10^{-4}$	–	–	$8,0 \times 10^{-3}$
Smelter	$4,2 \times 10^{-3}$	$1,1 \times 10^{-4}$	$9,4 \times 10^{-5}$	$1,7 \times 10^{-6}$	–	$4,4 \times 10^{-3}$
Caster	$3,7 \times 10^{-3}$	$4,4 \times 10^{-5}$	$7,8 \times 10^{-5}$	–	$5,9 \times 10^{-5}$	$3,9 \times 10^{-3}$
Blower	$3,7 \times 10^{-3}$	$4,4 \times 10^{-5}$	$7,9 \times 10^{-5}$	–	$5,9 \times 10^{-5}$	$3,9 \times 10^{-3}$
Crane driver	$3,2 \times 10^{-3}$	$4,4 \times 10^{-5}$	$7,6 \times 10^{-5}$	$5,9 \times 10^{-5}$	–	$3,4 \times 10^{-3}$
Cleaner	$4,2 \times 10^{-3}$	$1,5 \times 10^{-4}$	$9,4 \times 10^{-5}$	–	–	$4,4 \times 10^{-3}$
Fireproof materials handler	$3,4 \times 10^{-3}$	–	$7,6 \times 10^{-5}$	$1,7 \times 10^{-6}$	–	$3,5 \times 10^{-3}$
Foreman	$2,6 \times 10^{-3}$	–	$5,4 \times 10^{-5}$	–	–	$2,6 \times 10^{-3}$
Powder-gas catching device operator	$4,7 \times 10^{-3}$	$1,5 \times 10^{-4}$	$1,5 \times 10^{-4}$	–	–	$5,0 \times 10^{-3}$
Pumping unit operator	$3,7 \times 10^{-3}$	$4,4 \times 10^{-5}$	$7,3 \times 10^{-5}$	–	–	$3,8 \times 10^{-3}$
Electro-gas welder	$3,7 \times 10^{-3}$	–	$6,3 \times 10^{-5}$	–	–	$3,8 \times 10^{-3}$
Maintenance fitter	$5,1 \times 10^{-3}$	–	$7,5 \times 10^{-4}$	$1,7 \times 10^{-6}$	$5,9 \times 10^{-5}$	$5,9 \times 10^{-3}$
Electrician	$5,2 \times 10^{-2}$	–	$7,3 \times 10^{-5}$	$1,7 \times 10^{-6}$	$5,9 \times 10^{-5}$	$5,3 \times 10^{-2}$
Supervisory instruments and automation devices fitter	$4,7 \times 10^{-3}$	–	$7,5 \times 10^{-5}$	$1,7 \times 10^{-6}$	$5,9 \times 10^{-5}$	$4,8 \times 10^{-3}$

Allowing for the obtained CR values, we calculated the acceptable work period length at which the upper limit of acceptable occupational risk ( $10^{-3}$ ) was reached. As a result, average acceptable working period for workers employed at copper-smelting workshop amounted to 5 years.

Research on assessment of multi-environment population carcinogenic risks which were caused by exposure to carcinogenic substances from various

sources such as atmosphere, drinking water and food stuffs, showed that individual CR for population in a city where the examined enterprise was located amounted to  $2.3 \times 10^{-3}$  (the 4th risk range) [12]. The main contribution into multi-environment carcinogenic risk was made by arsenic, as it was in the case of occupational carcinogenic risk [11].

We determined Cyfra 21.1, CEA and NSE tumor markers in blood serum in the

course of periodical medical examination (PME) of workers. Our research revealed that 73% examined workers had excess levels of tumor markers; 19% workers had excess levels of two tumor markers at once. 9% workers had increased Cyfra 21.1. tumor marker; 14.5%, increased CEA tumor marker; 59%, increased NSE tumor marker.

The obtained data on carcinogenic danger of copper-smelting production were also confirmed by the results of epidemiologic study on mortality caused by malignant neoplasms among workers employed at copper-smelting workshop. Intensive mortality indices for male workers employed at smelting and converting sections of copper-smelting workshop amounted to 153.14 as per all localizations (totally). This parameter amounted to only

127.25 per 100,000 among ordinary male population. Intensive mortality indices for workers employed at blister copper production which were higher than the same indices for population were detected for malignant neoplasms of respiratory organs and chest (86.78 and 47.72 correspondingly), including lung cancer (71.47 and 43.48 correspondingly). The greatest difference in levels of lung cancer mortality among workers employed at copper-smelting workshop and control population was detected in 50-59 age group (425.53 and 159.57 correspondingly) ( $p < 0.05$ ).

As we can see from Table 2, statistically significant multiplicity of observed mortality excess over expected mortality was detected among men employed at copper-smelting

Table 2

Ratio of observed mortality caused by malignant neoplasms to expected mortality among men employed at copper-smelting workshop (per 100,000 people)

No.	Neoplasms localization	Observed	«Expected»	Observed indices and expected indices ratio
1	Oral cavity and pharynx	–	$3,70 \pm 0,88$	–
2	Respiratory organs and chest, including:	$86,78 \pm 21,04$	$40,70 \pm 2,94$	2,13*
	- trachea, bronchial tubes, lungs	$71,47 \pm 19,09$	$37,40 \pm 2,82$	1,91
	- nasal cavity and larynx	$15,31 \pm 8,84$	$2,52 \pm 0,73$	6,07
	- pleuroperitoneum and mediastinum	–	$0,78 \pm 0,41$	–
3	Digestive organs and peritoneum organs, including:	$25,52 \pm 11,41$	$43,92 \pm 3,05$	0,58
	- esophagus	–	$2,04 \pm 0,66$	–
	- stomach	$15,31 \pm 8,84$	$24,57 \pm 2,28$	0,62
	- bowels	$10,21 \pm 7,22$	$6,49 \pm 1,17$	1,57
	- liver	–	$3,48 \pm 0,86$	–
	- pancreas	–	$5,88 \pm 1,12$	–
	- others	–	$1,46 \pm 0,55$	–
4	Musculoskeletal system, connective tissue, skin, including:	–	$4,32 \pm 0,96$	–
	- skin	–	$1,12 \pm 0,49$	–
	- bones and connective tissue	–	$3,20 \pm 0,82$	–
5	Urogenital organs, including:	$15,31 \pm 8,84$	$8,90 \pm 1,37$	1,72
	- genital organs	$10,21 \pm 7,22$	$2,19 \pm 0,68$	4,66
	- urinary organs	$5,10 \pm 5,11$	$6,71 \pm 1,19$	0,76
6	Lymphatic organs and hematopoietic system	$5,11 \pm 5,11$	$5,63 \pm 1,09$	0,91
7	Others	$20,42 \pm 10,21$	$5,34 \pm 1,06$	3,82
8	All localizations taken together	$153,14 \pm 27,94$	$112,51 \pm 4,88$	1,36

Note: \* - discrepancies are statistically authentic ( $p < 0.05$ ).

workshop as per malignant neoplasms in respiratory organs and chest (2.13 times), including tumors in trachea, bronchial tubes and lungs (1.91 times), and nasal cavity and larynx (6.07 times). Besides, observed mortality exceeded expected mortality as per malignant neoplasms in urogenital organs (1.72 times), including genital organs (4.66 times), intestine tumors (1.57 times) and other localizations (3.82 times).

The obtained results will help to work out a system of CR managing which will include regulation and surveillance activities, organization and management activities, technical and technological activities, financial and economic activities, medical and prevention activities, rehabilitation, as well as information and educational activities. Providing hygienic safety and opportunity of technical achievement of goals set forth in risk managing are to become top criteria here; they should allow for economic efficiency of management decision-making. In future we should use technologies of insuring civil liability in relation to unforeseen damage to health and voluntary medical insurance of working population.

### Conclusions:

1. In blister copper production, carcinogenic risk caused by exposure to arsenic, cadmium, lead, nickel, beryllium, and benzpyrene, lies within unacceptable range and is determined mostly by arsenic as other basic carcinogens content doesn't exceed MPC.

2. The biggest carcinogenic risk values are detected for occupations with workplaces where dust emissions are the greatest, as well as for workers occupied with repair and auxiliary work.

3. Most workers who have occupations with unacceptable CR level also have increased levels of tumor markers.

4. Discrepancy between intensive mortality indices as per malignant neoplasms of different localization among workers employed at copper-smelting workshop and ordinary population proves there is an authentic correlation between neoplasms and impacts exerted by carcinogenic risks of production environment.

5. The obtained results will help to give grounds for a set of measures aimed at managing carcinogenic risks caused by shopfloor activity.

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