Δελτίο Ελληνικής Γεωλογικής Εταιρίας τομ. XXX/4, 57-64, 1994 Bulletin of the Geological Society of Greece vol. XXX/4, 57-64, 1994 Πρακτικά 7^{ου} Επιστημονικού Συνεδρίου, Θεσσαλονίκη, Μάιος 1994

THE PROBLEM OF SELENIUM DEFICIENCY IN SERBIA

Z. Maksimovic¹

ABSTRACT

Data on Se-deficiency in Serbia are presented. The results include Se contents of rocks, stream sediments, soil, cereal crops and garlic grown in the investigated soil, and human serum and scalp hair from several towns and regions. All data indicate a serious Se-deficiency. Analyses of human tissues show a very low Se-status of Serbian population. In some regions, Se contents of grains, garlic and human serum and scalp hair are approaching those of the low-selenium zone in China. It is assumed that the very low Sestatus of the human population could be a risk factor in the development of Balkan Endemic Nephropathy (BEN) and a high incidence of cancer of all sites in endemic areas.

INDRODUCTION

For mote than 35 years Se has been recognized as an essential nutrient to mammals and birds. Well-defined deficiency syndromes have been described in domestic animals and fowls. In humans, Se was found to have protective effects against a fatal cardiomiopathy, known as Keshan disease (Yang et al., 1984), and prophylactic and therapeutic effects against Kaschin-Beck disease, an endemic osteortopathy that occurs in the low-Se belt in China (Liang et al., 1986). Cancer, cardiovascular disease and muscular disorders were also discussed in connection with Se-deficiency (Selenium, WHO, Geneva, 1987, Scrauzer, 1992, Van Vleet et al., 1992). A hypothesis that very low Se-status of the human population in Serbia possibly plays an important role in the development of BEN was recently proposed (Maksimovic, 1987, 1991).

In this paper available data on Se-deficiency in Serbia are summarized, based on a team work during the last 10 years (Maksimovic et al., 1985, 1989, 1991, 1992).

ANALYTICAL PROCEDURE

In all samples, after wet digestion, Se was determined by hydride generation AAS. A Perkin Elmer atomic absorption spectrometer 5000, equipped with hydride generation system (MHS 10), was used. Different digestion procedures were used for geological and biological materials, and serious precautions were taken to prevent any loss of Se during this procedure (Maksimovic et al., 1991). The limit of detection was about 2 ng Se mL-1. Reproducibility of the technique was from -24 % to -2 %, depending on the concentration of Se. The accuracy of the analysis was monitored by inclusion of international reference samples in the analytical program (Maksimovic et al., 1992).

¹ Faculty of Mining and Geology, Djusina 7, 11000 BELGRADE, Serbia

SELENIUM IN STREAM SEDIMENTS

The first geochemical investigation of Se in Serbia, on a large scale, started on stream sediments (Maksimovic et al., 1985). The results obtained have shown that Se is more or less uniformly distributed in river sediments and the averages between different regions had small variations. The mean Se content of stream sediments, fraction <50 μ m, was 229 μ m/kg Se. It was much lower that the mean Se of soils in many countries (Thornton et al., 1983). The low Se content of the stream sediments, in spite of the contribution of selenium in numerous sulfide mineralizations, was the first indicator of Sedeficiency in the large part of the country.

SELENIUM IN IGNEOUS AND METAMORPHIC ROCKS

Selenium was determined in 285 igneous rock samples, including intrusive and volcanic rocks (Maksimovic et al., 1992). The overall average of 46 μ g/ kg Se is very close to the value of 50 μ g/kg Se proposed by Turekian and Wedepohl (1961) and Vinogradov (1962). Selenium content shows a general tendency to decrease from mafic to more acid rock types. Volcanic rocks are more depleted in Se than their intrusive and dike equivalents. The most widespread igneous rocks are those of intermediate composition. They are very poor in selenium, especially their volcanic equivalents, which are extremely depleted in this element (x=22 μ g/kg Se, n=90).

The average Se content in 89 samples of various metamorphic rocks is $28 \mu g/kg$ (Jovic et al., 1993). It was concluded that a great part of Se was lost from rocks during metamorphic processes at high temperature and pressure.

Very widespread volcanic rocks of intermediate composition and metamorphic rocks, both very depleted in Se, are the source of Se-deficient soil. Therefore, Se-deficiency partly originate from these Se-poor rocks, which built up vast volcanic and metamorphic areas in this country.

SELENIUM IN SOIL, WHEAT, CORN AND GARLIC

Samples of surface soil, and wheat, corn and garlic grown in these soil, were collected from several agricultural regions in Serbia, including plains in the north (Banat), river valleys of the Great Morava, West Morava, Drina and Kolubara rivers, as well as hilly areas of the west and central Serbia.

Soil

Se-content of soil derived from volcanic rocks of East Serbia is very low, $x=126 \mu g/kg$ Se (Jovic, 1990). All other investigated soils were derived from sedimentary rocks of different age and composition (from quaternary to Paleozonic). They are all poor in selenium. The mean content of Se in 140 soil samples from 29 communities in Serbia is 200 $\mu g/kg$, a value about twice lower than those found in soil from other regions of the world (Thornton et al., 1983). The range of selenium in the soil, 79 to 439 $\mu g/kg$ Se, shows that the highest value found in the soil is less than 500 $\mu g/kg$, which is considered as upper limit for Se-deficient soils (Mayland et al., 1989).

Results for soils, grain and garlic are summarized in Table 1.

PH of soil varies from 4.45 to 8.33, the acidic to neutral soils being the most frequent. These conditions promote low availability of selenium in plants. Se extractable with $1N NH_4Oac$ varies from 1.2 to 27.6 % of the total, depending mostly on pH of the soil. The highest pH of soil was found in the areas of south Banat in the northern Serbia, where amounts of extractable selenium was the highest.

	n	Mean ± SD µg/kg
Soils	140	200 ± 69.6
Wheat	58	20.5 ± 12.4
Corn	79	13.7 ± 13.6
Garlic	66	13.7 ± 17.1

Table 1: Se content in Serbian soils, wheat, corn and garlic

n=number of samples; wheat, grain and corn of dry wt basis and garlic fresh after peeling.

Wheat

Se-content of wheat from Serbia is low and ranges from 3.6 to 65.5 µg/kg with the mean value of 20.5 µg/kg Se from 58 samples. There are wide variations of the Se-content of wheat in various regions. The highest concentration was found in two communities in the central Serbia (Zabari and Svilajnac) (\bar{x} =39.2 µg/kg; n=3) and in the Banat areas, north Serbia (x=25.2 µg/kg; n=31). The lowest value of selenium originate from hilly areas of the west Serbia (\bar{x} =9.4 µg/kg; n=14), where pH of the soil is acidic. It should be noted that Kechan disease in China affects children and young women in areas where the mean level of Se in staple grains is less than 25 µg/kg (Diplock, 1986).

Corn

Corn is very poor in selenium, in the range of 2.8 to 82.0 μ g/kg, with the average of 13.7 μ g/kg Se from 79 samples. Two communities in the central Serbia (Zabari and Svilajnac) have the highest Se-content in corn (\bar{x} =58.9 μ g/kg; n=3). However, most of the communities have extremely low Se levels in corn (<10 μ g/kg). These values correspond to those found in corn in the low-Se belt in China (Xu and Jiang, 1986). They also correspond to values found in corn from Pozega Valley in Slavonija, Croatia, where mass occurrences of Se responsive diseases in domestic animals are connected with Se deficiency in soil and fodder (Gavriloc and Matesic, 1986).

Garlic

Garlic is known to accumulate more selenium than most of other crops. The highest content of 81.40 μ g/kg Se was found in garlic from seleniferous soil in the Enshi County, China, where chronic selenosis in human population was recognized (Yang et al., 1983). Reported Se content in garlic from U.S.A. was 250 μ g/kg (Morris and Levander, 1970) and 600 μ g/kg (Mikkelsen et al., 1989). Compared with these levels in the U.S.A. one reported value from United Kingdom of 20 μ g/kg is very low (Thorn et al., 1978).

Garlic is very popular vegetable in Serbia and is grown practically in every garden. Garlic and soil were collected from 66 sites in 21 communities. The analyses revealed wide variations of Se level in garlic from different regions, with the mean content of 13.7 μ g/kg Se,. The highest content was found in three communities in the north Serbia (Banat), with average of 35.4 μ g/kg Se from 13 sites. In these cases total and extractable selenium was relativelyhigh and the soil was alkaline. Two communities with a moderate Se content in wheat and corn in the central Serbia (Zabari and Svilajnac) have higher Se content than the overall average in Serbia (x=18.9 μ g/kg; n=5). However, most of the communities have extremely low Se levels in garlic (<10 μ g/kg), not reported in the literature so far.

SELENIUM IN HUMAN SERUM AND SCALP HAIR

In recent years it has been establishes beyond doubt that selenium is also essential to humans and diseases and death resulting from Se-deficiency have been described (Chen et al., 1980; Xu and Jiang, 1986).

In order to examine the levels of selenium in serum and scalp hair of Serbian population and subsequently to correlate these with the incidence of degenerative diseases in the respective areas, a collaborative study started in 1988 under the auspice of Serbian Academy of Sciences and Arts in Belgrade.

The samples of human serum and scalp hair were obtained from healthy individuals, aged 20-50, from several towns and regions, but not including villages with BEN.

In this country interchange of foodstuffs occurs from one district to another. However, low selenium content in soil and grain in the main agricultural regions is reflected in the low Se-status of human population. Serum Se-levels for a combined male and female population from the individual regions in Serbia are summarized in Table 2.

Presented data indicate a Se deficiency in human population in Serbia.

n	Mean Se	±SD µg/I						
109	45.0	16.2						
303	47.7	24.7						
15	47.0	18.2						
18	32.1	19.5						
59	38.3	17.7						
98	37.9	20.9						
602	44.2	±19.5						
	109 303 15 18 59 98	109 45.0 303 47.7 15 47.0 18 32.1 59 38.3 98 37.9						

Table 2: Mean serum Se-level in healthy population aged 20-50 from various parts of Serbia

Serum Se-levels are lower than in any country in Europe (Thorling et al., 1986). Epidemiological investigation in Finland has demonstrated an increased risk of cancer in people with serum Se-levels at 45 μ g/L and below (Salonen et al., 1985). This border line deficiency may be present, according to Thorling et al., 1986, in part of the Greek population. In Serbia the areas with the mean Se-levels below 45 μ g/L are quite common (Fig. 1).

High correlation (r=0.85) was found between Se-levels in serum and scalp hair. The Se-content in hair is very low in all investigated areas, approaching those in the Se-deficient zone in China.

Ta	ble	3:	Mean	Se	levels	in	scaip	hair	of	helathy	population	aged	20-50
from	var	ious	parts	of	Serbia								

Location	n	Mean Se	±SD µg/L
Vojvodina (north Serbia)	72	91	21
Belgrade area	246	96	17
Central & West Serbia	12	109	14
East Serbia	44	89	22
South Serbia	6	88	14
Serbia	380	94	±16

Ψηφιακή Βιβλιοθήκη "Θεόφραστος" - Τμήμα Γεωλογίας. Α.Π.Θ.

60



Fig. 1: Mean serum Se-levels of healthy population from Serbia, compared with those from some other countries.

All those results demonstrate that low Se-status of Serbian population correlates with low Se content of food supplies.

Se DEFICIENCY AND POSSIBLE EFFECT ON HEALTH

There is no record in Serbia of specific disease in humans caused by Se deficiency. However, series Se deficiency could be one of the main etiological agents of BEN and high incidence of Urinary Tract Tumors (UTT) in endemic areas (Maksimovic. 1987; Maksimovic et al., 1989; Maksimovic, 1991). Geographic correlation between BEN and the highest incidence of UTT were established in Yugoslavia and Bulgaria (Petrovic, 1968; Chernozemsky et al., 1977). This high percentage of UTT in patients and regions with BENN is indicative of a common etiological agent that has nephropathogenic and carcinogenic effects.

So far, Se deficiency is the only observed common demoninator for all endemic areas. Taking into account the biological function of Se (Diplock, 1981), an underlying significant Se deficiency may predispose individuals to potentially nephrotoxic and carcinogenic agents. These data suggest that research of BEN and UTT in endemic areas is a critical avenue to explore.

Results of current research on Se levels in patients from Serbia with cancer (276 subjects) and myocardial ischemia (20 subjects) show that these patients have lower Se levels for about 20 % compared to controls (Maksimovic et al., 1992). Based on these data and those reported for

population groups in other countries (Combs and Combs, 1986), it appears that Se is a possible dietary inhibitor of cancer and myocardial ishemia promotion.

REFERENCES

- CHEN, X., G., CHEN, J., CHEN, X., WEN, Z. and GE, K. (1980). Studies on the relation of selenium and Keshan disease Biol. Trace Elem. Res. 2, 91-107.
- CHERNOZEMSKY, I.N., STOYANOV, I.S., PETKOVA-BOCHAROVA, T.K., NIKOLOV, I.G., DRAGANOV, I.V., STOICHEV, I.I., TANCHEV, Y., NAIDENOV, D. and KALCHEVA, N.D. (1977). Geographic correlation between the occurrence of endemic nephropathy and urinary tract tumors in Vratza district, Bulagria, Int. J. Cancer, 19, 1-11.
- COMBS, F.G., Jr. and COMBS, S.B. (1986). The role of selenium in nutrition. -Academic Press, Inc., Orlando, San Diego, New York.
- DIPLOCK, A.T. (1981). Metabolic and functional defects in selenium deficiency. -Phil. Tans. R. Soc. London, B294, 105-117.
- DIPLOCK, A.T. (1986). Selenium and selenium-responsive disease. In, I. Thornton (ed.) Environmental geochemistry and health. -Sci. Reviews Ltd, Norhwood, England, 181-191.
- GAVRILOVIC, B. and MATSIC, D. (1986). Importance of selenium quantity in soil and fodder in regard to the occurrence of some diseases in catle, pigs, sheep, and poultry in Yugoslavia. In, Proceed. 3rd Int. Symp. On Selenium in Biology and Medicine, Combs, G.F., Jr., Spallholz, J.E., Levander & Oldfield (eds)., Acvi Publi. Co., Westport, Conn., 740-749.
- GONDI, F., PANTO, GY., FEHER, J., BOGYE, G. and ALFTHAM, G. (1992). Selenium in Hungary. The rock-soil-human system. - Biol. Trace Elem. Res., 35, 299-305.
- JOVIC, V. (1989). Geochemistry of bioessencial trace elements in the processes of weathering of the Cretaceous-tertiary volcanic rocks in Serbia.-Coctor's Dissertation, Faculty of Mining and Geology, Belgrade, 380 p. (in Serbian).
- JOVIC, V., MILIC, S. and POPADIC, D. (1993). Selenium in some metamorphic rocks in Serbia.-Conference on Selenium, Belgrade 1993, Abstracts, p. 46-47.
- LIANG, S.T., ZHANG, J.C., SHANG, X., MU, S.Z. and ZHANG, F.J. (1986). Effects of selenium supplementation in prevention and treatment of Kaschin-Beck disease. -In, Proseed. 3rd Int. Symp. On Selenium in Biology and Medicine, Combs, G.F.Jr., Spallholz, J.E., Levander, O.A., & Oldfield, J.E. (eds)., Avi Publ. Co., Westport, Conn., 938-946.
- MAKSIMOVIC, Z. (1987). Trace element deficiency and Balkan Endemic Nephropathy (BEN). - In, Etiology of Endemic (Balkan) Nephropathy, Strahinjic, S. & Stefanovic, V (eds.), Nis, 43-47.
- MAKSIMOVIC, Z. (1991). Selenium deficiency and Balkan Endemic Nephropathy (BEN). Kidney International, 40. Suppl. 34, pp. S-12 S-14.

MAKSIMOVIC, Z., DANGIC, A., JOVIC, V. and RSUMOVIC, M. (1992). Selenium in the igneous rocks of Serbia (Yugoslavia). Bull. Acad. Serbe Sci. Arts, 33, 23-37.

MAKSIMOVIC, Z., DUJIC, I. and JOVIC, V. (1989). Deficiency of selenium in

Ψηφιακή Βιβλιοθήκη "Θεόφραστος" - Τμήμα Γεωλογίας. Α.Π.Θ.

62

the environment in the Eastern Serbia (region of Zajecar) and possible consequences to health. -Manand environment, 4-5, 24-32, Belgrade (in Serbian with English summary).

- MAKSIMOVIC, Z., DUJIC, I, I. and JOVIC, V. (1991). Selenium status of soil, grain and human population in Yugoslavia. -In, Trace Elements in Man and Animals, No. 7, 22-5 to 22-6, Momcilovic, B. (ed,), Zagreb, IMI.
- MAKSIMOVIC, Z., , DUJIC, I., JOVIC, V. and RSUMOVIC, M. (1992). Selenium dificiency in Serbia and possible effects on health. Bull. Acad. Serbe Sci. Arts, 33, 65-83.
- MAKSIMOVIC, Z., RSUMOVIC, M. and RADOSEVIC, P. (1985). Selenium in certain river sediments in Serbia (Yugoslavia). -Bull. Acad. Serbe Sci. Arts, 26, 103-109.
- MAKSIMOVIC, Z., NIKOLIC, M., JORGA, J., RSUMOVIC, M., RADOSEVIC, P., DACIC, V. and CUSIC, S. (1993). Selenim status of soil, grain, garlic and human serum and incidence of cancer in the community Barajevo (Belgrade). -Conference on Selenium, Belgrade 1993, Abstracts, p. 46-47.
- MAYLAND, H. F., JAMES, L. F., PANTER, K. E. and SONDEREGGER, J. L. (1989). Selenium in seleniferous environment. In, Selenium in agriculture and the environment, Jacobs, L.W. (ed), Spec. Publ. 23, 15-20, Madison, Wisconsin.
- MIKKELSEN, R.L., PAGE, A.L. and BINGHAM, F.T. (1989). Factors affecting selenium accumulation by agricultural crops. -In, Selenium in agriculture and the environment, Jacobs, L.W. (edit.), SSSA Special pubi., 23, 65-94, Madison Wisconsin.
- MORRIS, Y.C. and LEVANDER, O.A. (1970). Selenium content of foods. J. Nutrit. 100, 1383-1388.
- PETKOVIC, S. (1968). Geographical distribution of cancer of the urothelium in Yugoslavia. -Int. J. Urol. Stud. 35, 1-9.
- SALONEN, J.T., SALONEN, R., LAPPETELAINEN, R., MEANRAA, P.H., ALFTHAN, G. and PUSKA, P. (1985). Risk of cancer in relation to serum concentration of selenium and vitamin A and E: matched case-control analysis of prospective data. -Brit. Med. J. 290, 417-420.
- SELENIUM, Environmental Health Criteria 58, WHO, Geneva, 1987, pp. 198-237.
- THOMSON, C.D., REA, H.M., DOESBURG, V.M. and ROBINSON, M.F. (1977). Selenium concentrations and glutathione peroxidase activities in whole blood of New Zealand residents. -Br. J. Nutr. 37, 579-587.
- THORLING, E. B., OVERVAD, K. and GEBOERS, J. (1986). Selenium status in Europe - human data, a multicentar study. - Annalsof Clinical Res., 18, 3-7.
- THORN, J., ROBERTSON, J. and O'CONNOR, R. (1978). Trace nutrients. Selenium in British food. -Brit. J. Nutr. 39, 391-396.
- THORTON, I., KINNIBURGH, D.G., PULLEN, G. and SMITH, C.A. (1983). Geochemical aspects of selenium in British soils and implications to animal health. -In, Trace substances in environmental helath XVII, Hemphil, D.D. (ed.)., 391-398.
- TUREKIAN, K.K. and WEDEPOHL, K.H. (1961). Distribution of the elements in some major units of the crust of the Earth. -Geokhimia, 555-571.
- VINOGRADOV, A.P. (1962). Average content of chemical elements in the chief

types of igneous rocks of the crust of the Earth. -Geokhimia, 555-571. XU, G.L. and JIANG, Y.F. (1986) - Selenium and prevalence of Kechan and Kachin-Beck disease in China. -Proceed. 1st Int. Dymp. Geochem. Health, Thornton, I. (ed.), London, 192-204.

- YANG, G., GHEN, J., WEN, Z., GE, K., ZHU, L. and CHEN, X. (1984). The role of selenium in Keshan disease. - In, Advance in nutritional research, Draper, H.H. (ed.), Plenum Press, New York, 203-231.
- YANG, G., WANG, SH., ZHOU, R. and SUN. SH. (1983). Endemic selenium intoxication of humans in China. -Am, J. Clin. Nutr. 37, 872-881.
- ZACHARA, B.A. (1989). Selenium variables in human blood in some European countries. -3rd Int. Congress on Trace elem. In Health and Disease, Adana, Turkey, Abstract Book, p. 23.