

Endocrine Diseases in Post-Chernobyl Period in Belarus

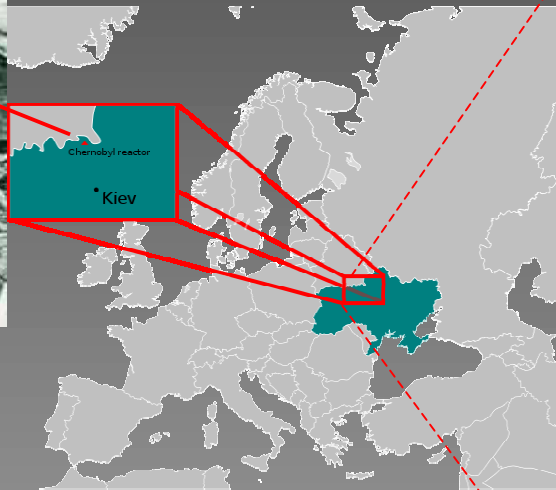
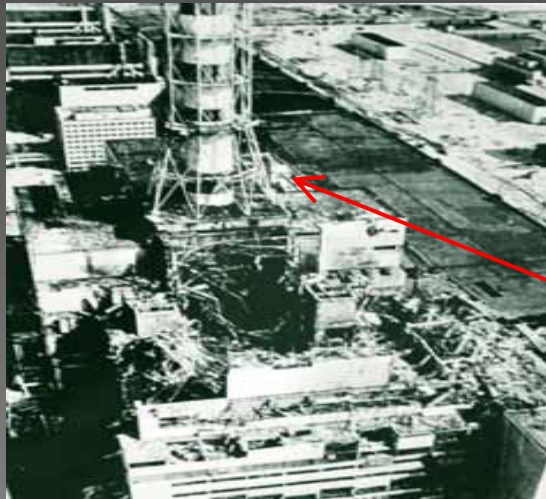
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26 April 1986

The accident at the Chernobyl nuclear power plant



Radioactive substance release - 13 EBq:

- ^{131}I - 1.8 Ebq
- ^{137}Cs - 0.085 Ebq
- ^{90}Sr - 0.01 Ebq
- Pu isotopes - 0.003 EBq

More than 200 000 sq km of Europe were contaminated with $> 37 \text{ kBq/sq m}$ of ^{137}Cs

Over 70 % of this area was in the three most affected countries, Belarus, Russia and Ukraine.

The Chernobyl Forum: 2003-2005, IAEA

- The accident at the No. 4 reactor of the Chernobyl Power Station took place on April 26, 1986, at 1:23 AM.
- Two explosions occurred, the first due to steam and a second one due to hydrogen. The explosions expelled fission products and fuel elements to the exterior that accumulated in a cloud reaching to approximately 7,000 m and centered at approximately 4,500 m.
- Because the graphite ignited, there was a second, more prolonged but less intense, release over a 9- to 10 day period that peaked on May 6, 1986, and dropped sharply on May 11 as the fire was extinguished. The following volatile elements, as well as the noble gases xenon and krypton, constituted the most abundant released material: Iodine, cesium, and tellurium.



INTRODUCTION

- Republic of Belarus belongs to European countries with predominantly light and moderate chronic iodine deficiency
- The State Program of iodine prophylaxis with iodinated salt or KI tablets was stopped in the end of 70-s yrs. and restored - during 1998-2000 yrs. (12 yrs. after the Chernobyl catastrophe);
- *There were no organized prophylaxis with stable iodine among the population during first hours/days/ weeks of the Chernobyl catastrophe;*

- Starting a week after the Accident and then - during all summer months in 1986yr. there was spontaneous individual intake of iodine solutions in different doses by different age group of population
- Side effects:
 - *Rashes*
 - *very firm thyroid gland, autoimmune thyroiditis*
 - *Wolf-Chaickoff effects*

Iodide Prophylaxis in Poland After the Chernobyl Reactor Accident: Benefits and Risks

J. NAUMAN, J. WOLFF//American J Medicine.1992 - V.94: 524-532.

In Poland, decided to adopt the very conservative dose commitment - increased air radioactivity was first detected on the of 50 mSv as the intervention level for children.

The commission on the morning of April 29: Whole body committed dose should not exceed 5 mSv/y (0.5 rem). Thyroid committed dose should not exceed 50 and amounted to 504 Bq of radioiodines /m³ air with 1.55- to 3.0~times higher values in northeast Poland. Approximately 18 million doses of KI solution were sources distributed;

“95.3% of children received iodide prophylaxis;

86.7% took a single dose,

2.39% took two or more doses,

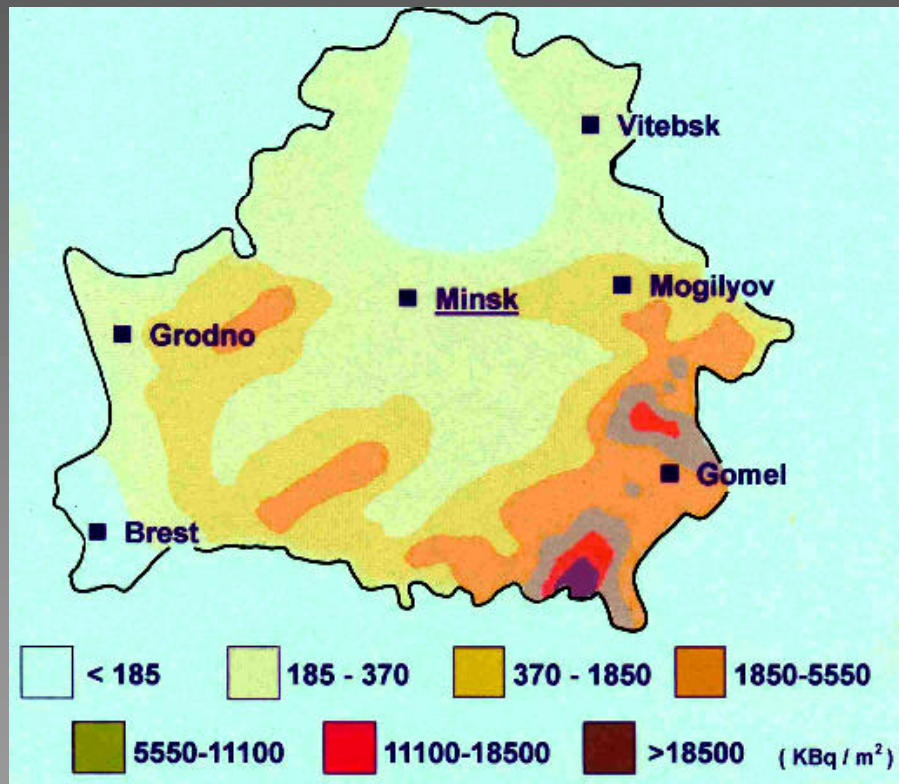
and a surprising 6.14% were given diluted tincture of iodine by their parents before the start of the program and then took a single dose of KI”

*Approximately 10.5 million children 16 years old and under,
and approximately 7 million adults received iodide
prophylaxis in Poland*

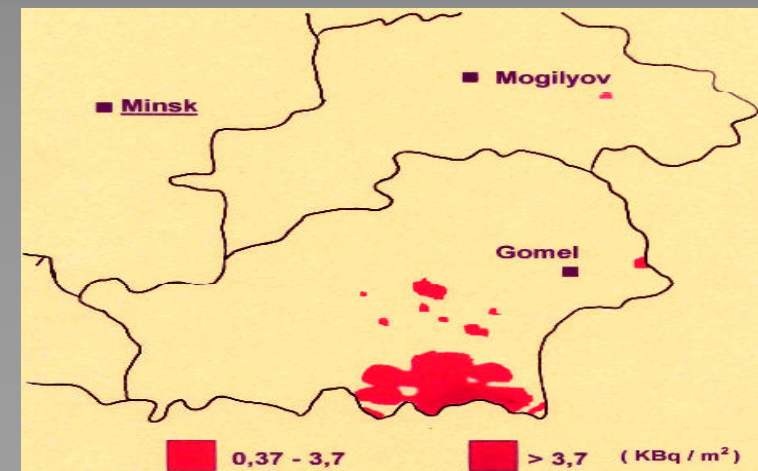
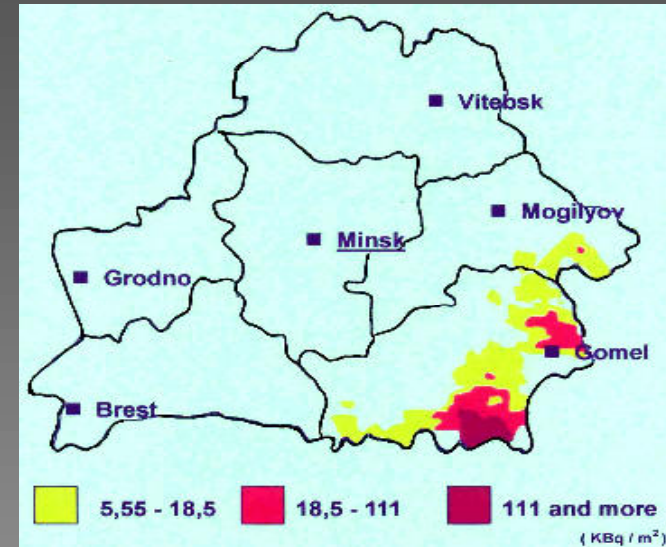
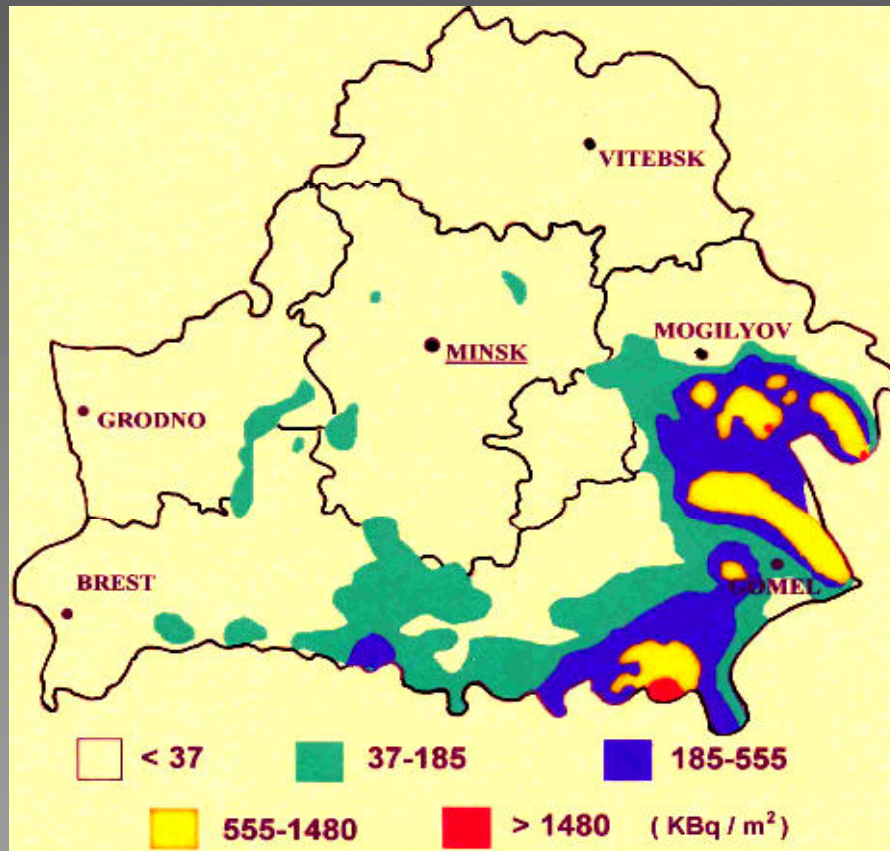
*IODIDE PROPHYLAXIS IN POLAND AFTER
THE CHERNOBYL REACTOR ACCIDENT:
BENEFITS AND RISKS*

*J. NAUMAN, J. WOLFF//AMERICAN J MEDICINE.1992 - V.94: 524-
532.*

Distribution of Iodine-131 in the Soil (May 10, 1986)



Belarusian Territory: contamination with Cesium-137, Strontium-90 and Plutonium-238, 239, 240 (January 1, 1995)



Effect of chronic iodine deficiency and stable iodine consumption

Table 4. Estimated risk of developing thyroid cancer after a radiation dose of 1 Gy, by level of soil iodine in the settlement of residence at the time of the accident and by potassium iodide (i.e., antistrumin) consumption status (analyses restricted to subjects with radiation doses to the thyroid of less than 2 Gy)*

Consumption of potassium iodide	OR at 1 Gy (95% CI)	
	Highest two tertiles of soil iodine	Lowest tertile of soil iodine
No	3.5 (1.8 to 7.0)	10.8 (5.6 to 20.8)
Yes	1.1 (0.3 to 3.6)	3.3 (1.0 to 10.6)

*Levels of iodine in soil in settlement of residence at time of accident were divided into tertiles. OR = odds ratio at 1 Gy compared with no exposure; CI = confidence interval.

Exposure to ionizing radiation is the only established risk factor of thyroid cancer in early childhood

Cardis E, Kesminiene A, Ivanov V, Malakhova I, Shibata Y, Khrouch V, et al. Risk of thyroid cancer after exposure to ¹³¹I in childhood. J Natl Cancer Inst 2005;97:724–32.

Tab. 2. Excess relative risk for thyroid carcinoma. Comparing Chernobyl to external radiation, and comparing the effect of iodine intake on the risk (5)

	Chernobyl overall	External radiation	Chernobyl high iodine	Chernobyl low iodine	Chernobyl high I + KI	Chernobyl low I + KI
ERR	4,5-7,4	7,7	2,5	9,8	0,1	2,3

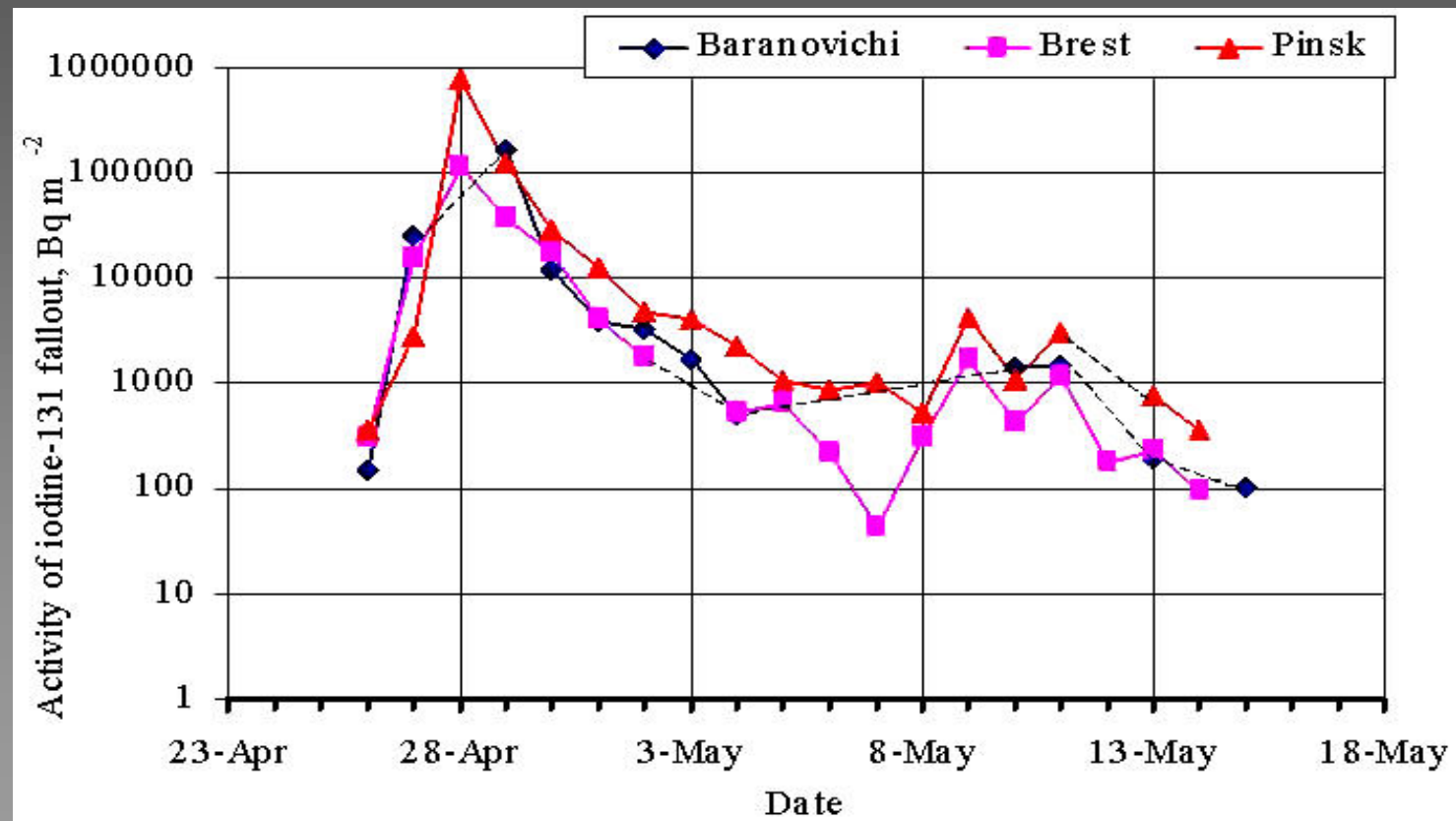
Screening for Chernobyl-associated disorders in Belarusian population started in May 1986



Screening data of thyroid nodules and thyroid cancer in Belarus after the Chernobyl accident

Screening programs	Year of screening	N of subjects	Diseases			
			Thyroid nodes		Thyroid carcinoma	
			n	%	n	%
ICP, IAEA (Mettler et al, 1992)	1990	323	4	1,24	-	-
Screening program of Research Institute of Radiation Medicine, Choiniki, Gomel oblast, Belarus (Drozd et al., 1993, Drozd et al., 2002)	1990-1991	1132	14	1,24	7	0,62
	1993	1546	32	2,1	5	0,3
IPHECA, WHO (Technical repot,1996)	1993-1994	14000	210	1,5	32	0,22
Chernobyl Sasakawa Project. Sasakawa Memorial Health Foundation (Yamashita, 1999) Mogilev oblast Gomel oblast	1990-2000					
		13868 19790	24 350	0,1 1,7	2 38	0,008 0,19
A Cohort study of Thyroid cancer and thyroid diseases after the Chernobyl Accident. Chernobyl disease study group of Belarus and USA Ostapenko et al.,2001 Stezhko V. et al. 2004	1996-2001 2004					
		11200 25 161	694 n/I	6,2 n/i	53 100	0,47 0,4
Screening project of Red Cross (Brest oblast) (unpublished data)	1998-2008	164175	23693	14,4	499	0,3

Dynamics of the measured daily Deposition Density of ^{131}I in the Cities of Brest Oblast



Yu. Gavrilin, S. Shinkarev, A. Bouville, M. Germenchuk, M. Hoshi, N. Luckyanov, P. Voilleque, O. Zhukova «Retrospective Assessment Of Thyroid Doses For The Residents Of Brest Oblast Of Belarus», 2004

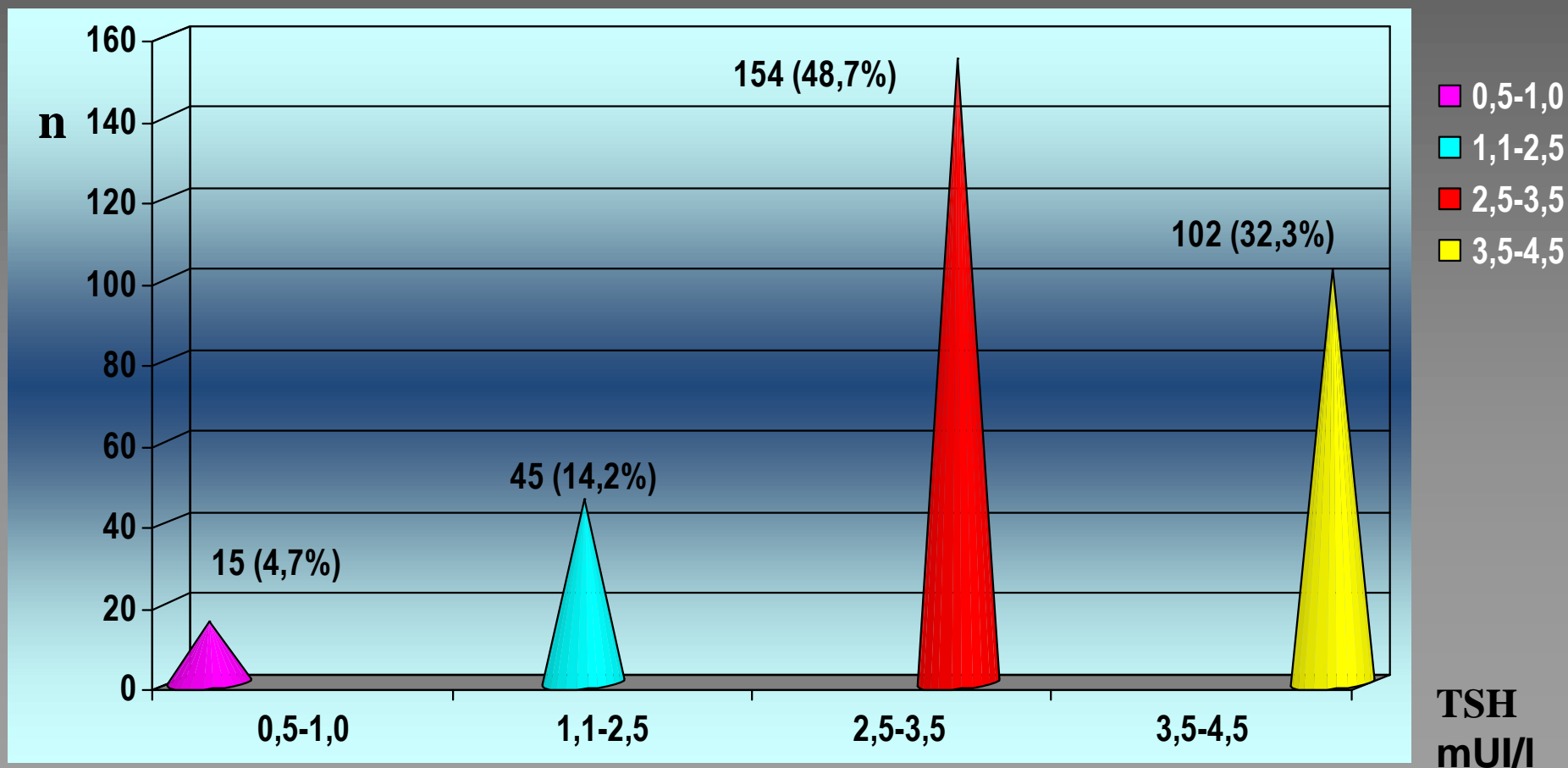
Screening in Stolin Brest region (1996-2008)



- In 2000-2011 active screening for thyroid diseases was performed in women of reproductive age in Minsk and Brest regions
 - Percentage of women with individual levels of TSH between 2.5-3.5 mIU/l was 48.7%, 3.5-4.5 mIU/l – 9.4%, > 4.5 mIU/l – 8.2%
 - Ultrasound abnormalities of thyroid gland were revealed in 44.6% examined women(among them – 23.4% were TPO-Ab positive)
- 2000-2002 fT4 and TSH-screening in pregnant women revealed non optimal distribution of individual levels during first 9 weeks of pregnancy
- Next fT4 and TSH-screening was performed in pregnant women (9 weeks of pregnancy) in 2009-2011 yrs

Distribution of the cases of different TSH levels in women during first 9 weeks of pregnancy (2000-2002)

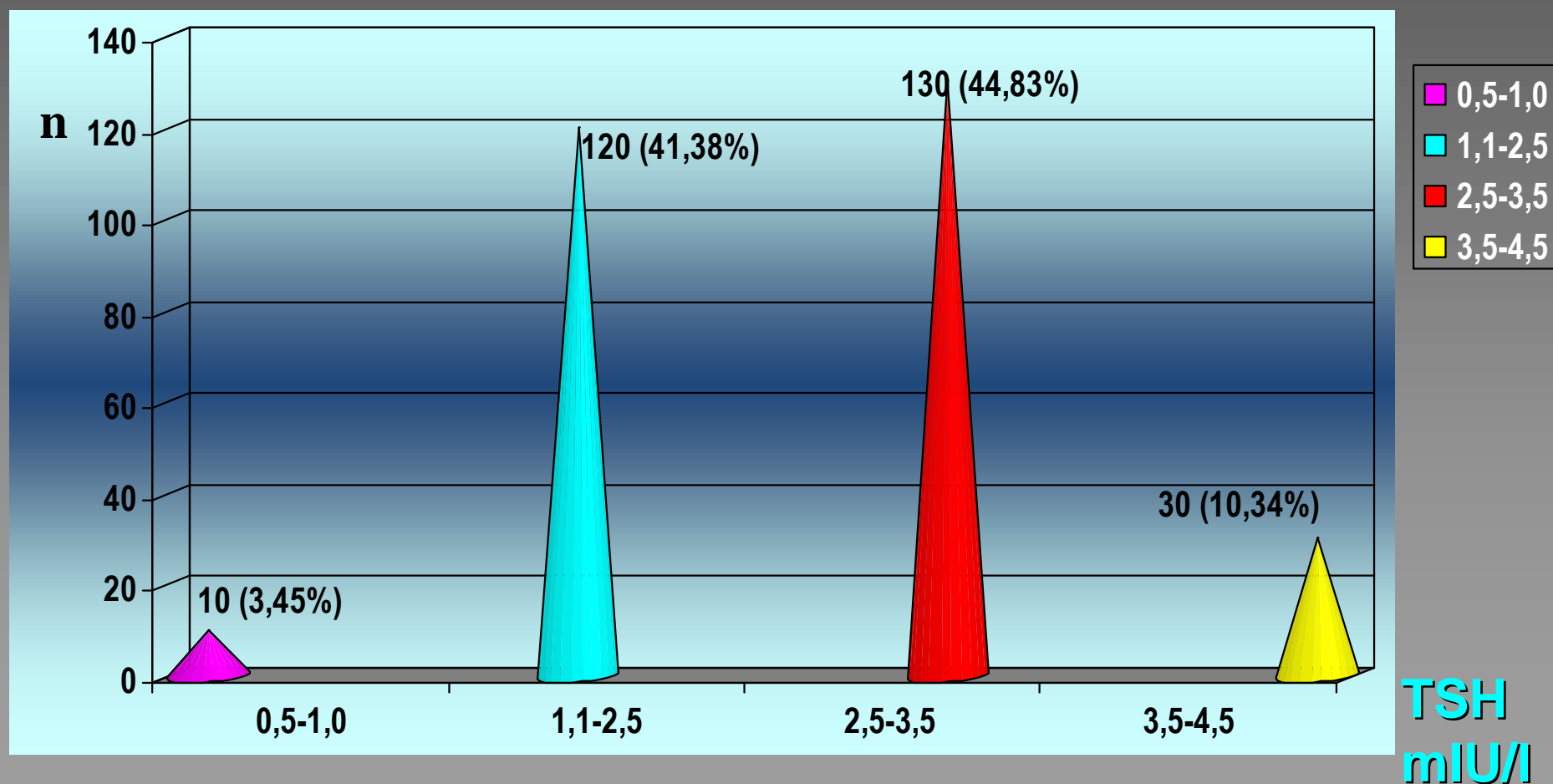
N=316



E.M.Kapustina et al., 2002.

Distribution of the cases of different TSH levels in women during first 9 weeks of gravities (2009-2011)

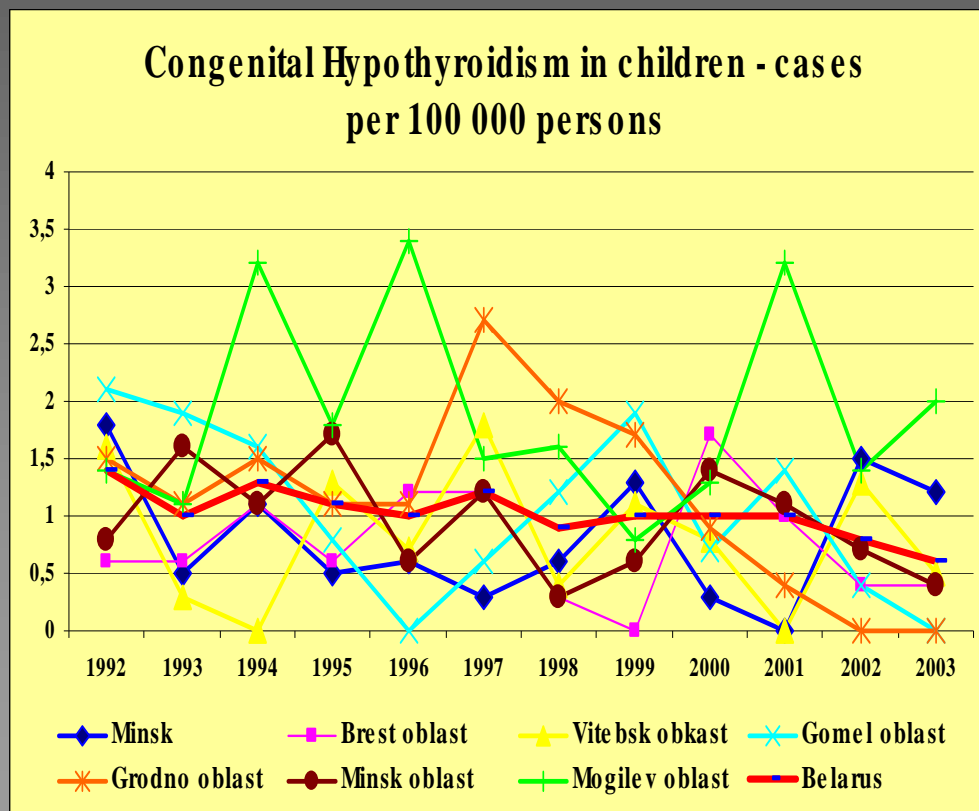
N=290



Iodine excretion screening in Brest and Minsk regions

- According to results of screening in some districts of Brest and Minsk regions - the situation is changing to the best.
 - The median levels of iodine excretion in Stolin (Brest region) in 2002 yr were 36.4 $\mu\text{g/L}$ (18.3 / 47.5) in adults and 43.6 $\mu\text{g/L}$ in schoolchildren.
 - In Minsk region median levels of iodine excretion during the same period were 59.8 $\mu\text{g/L}$ in adults and 75.4 $\mu\text{g/L}$ - in schoolchildren.
- In 2007 yr the median level of iodine excretion in children of
 - Stolin reached 76.3 $\mu\text{g/L}$
 - Minsk region - 73.2 $\mu\text{g/L}$.

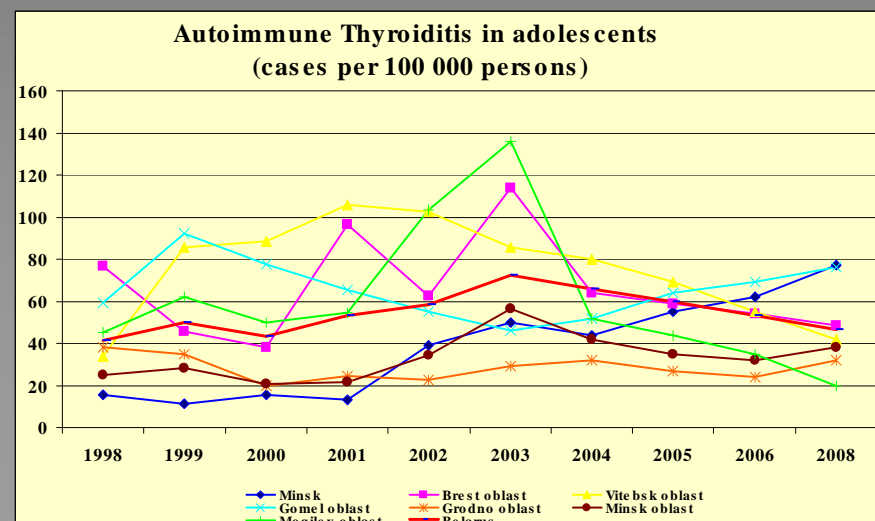
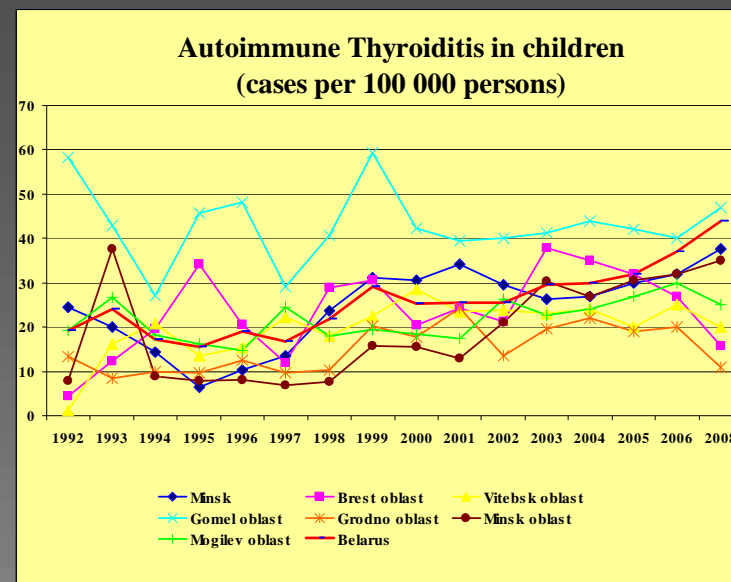
Congenital Hypothyroidism



Screening programs for
congenital hypothyroidism
Thyroid tests and
Prophylaxis with stable
iodine during pregnancy

Autoimmune thyroiditis

- Increased incidence of autoimmune endocrine diseases in children and adolescents (Autoimmune thyroiditis during 1995-2000 yrs.)
- Clinical features of autoimmune thyroiditis in groups of patients under 25 yrs. – subclinical forms and coexistence with nodular pathology, coexistence with DM type 1;
- Retardation/ impairments of physical and sexual development in children and adolescents



Possible Etiologic Factors of Thyroid Diseases

Endogenous factors

- Gender
- Age
- Ethnic factors
- Hormones
- Pregnancy

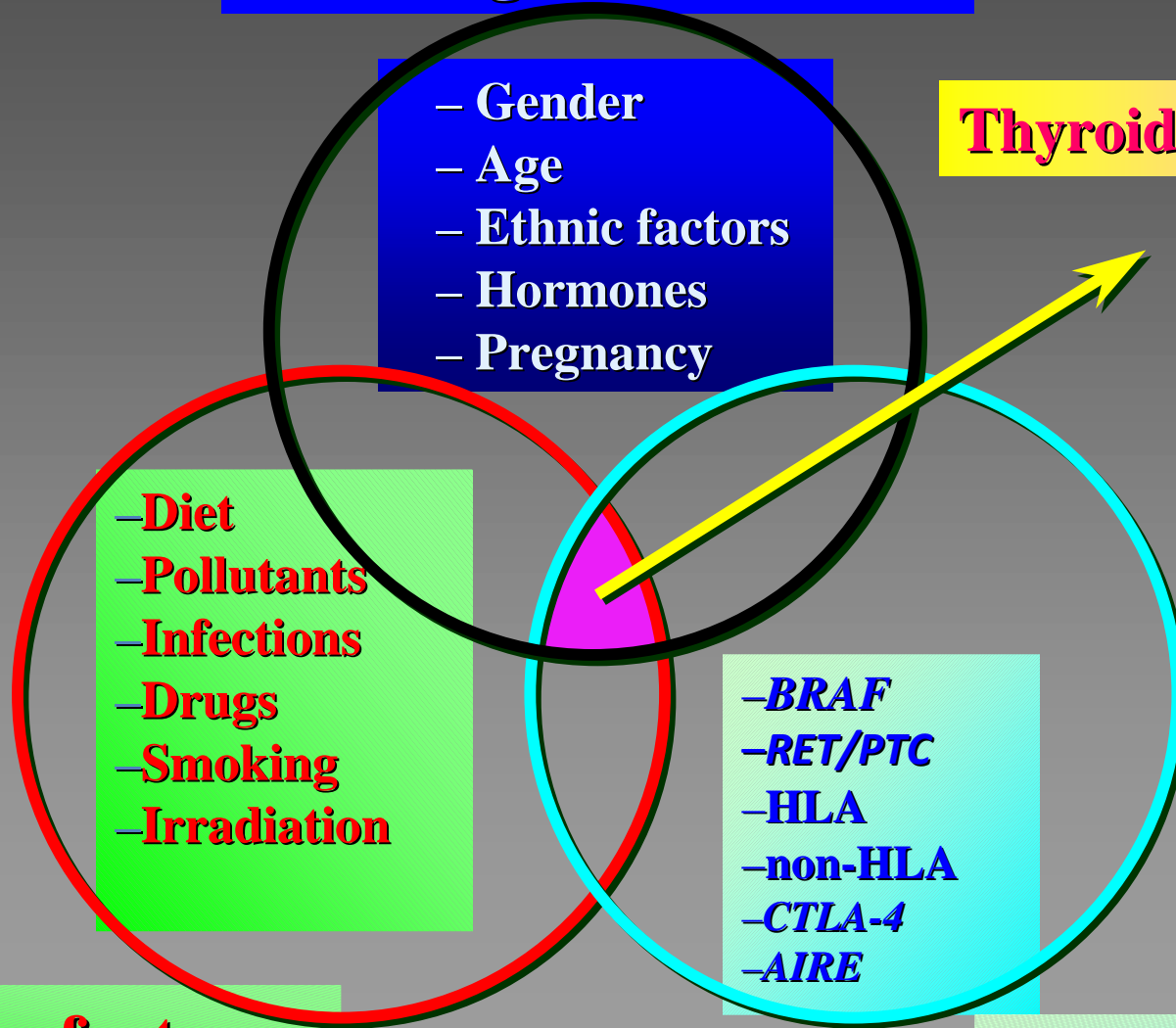
Thyroid Diseases

- Diet
- Pollutants
- Infections
- Drugs
- Smoking
- Irradiation

Exogenous factors

- *BRAF*
- *RET/PTC*
- HLA
- non-HLA
- *CTLA-4*
- *AIRE*

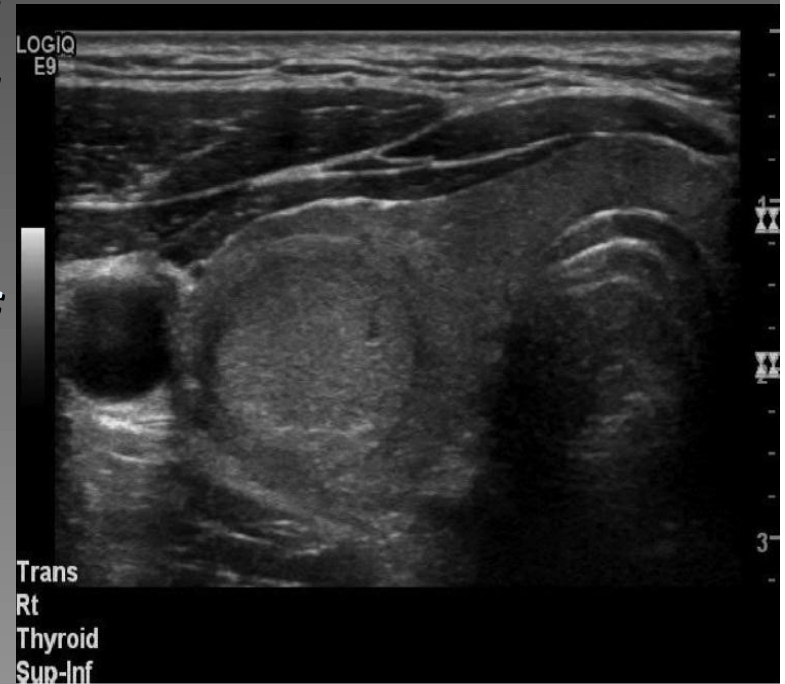
Genetic factors



Thyroid nodules in clinical practice

Their prevalence depends to a great extent on the method used for detection. With the increased utilization of US for evaluation of the lesions of the neck, the incidental finding of unsuspected thyroid nodules has dramatically increased. The prevalence of thyroid nodules is higher in women in areas of iodine deficiency and increases with advancing age.

Factors that increase the risk of malignancy include radiation exposure, history of head and neck irradiation, and very young or advanced age

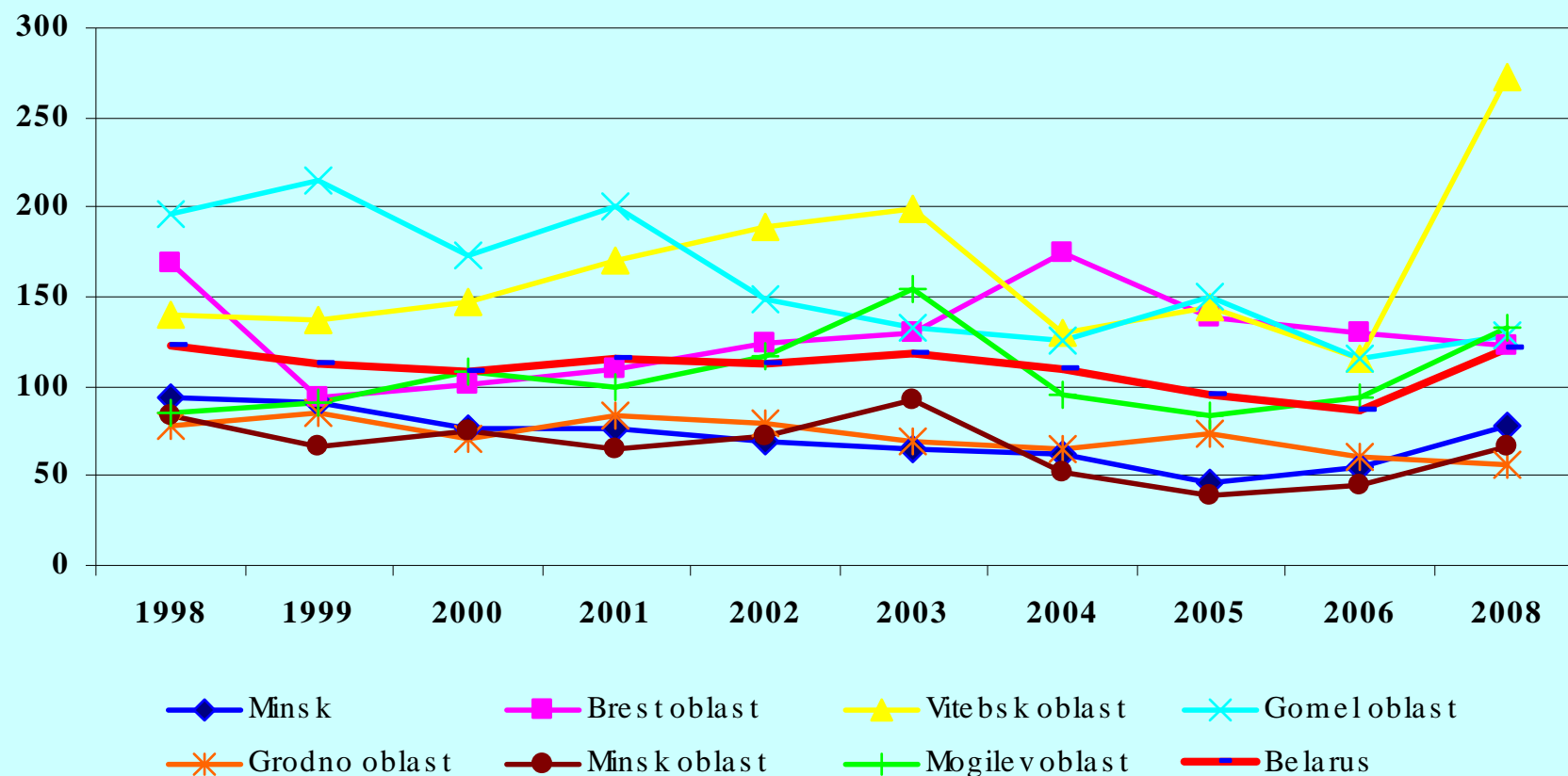


TR.S. Bahn, M. R. Castro / J Clin Endocrinol Metab, 2011, 96(5):1202–1212

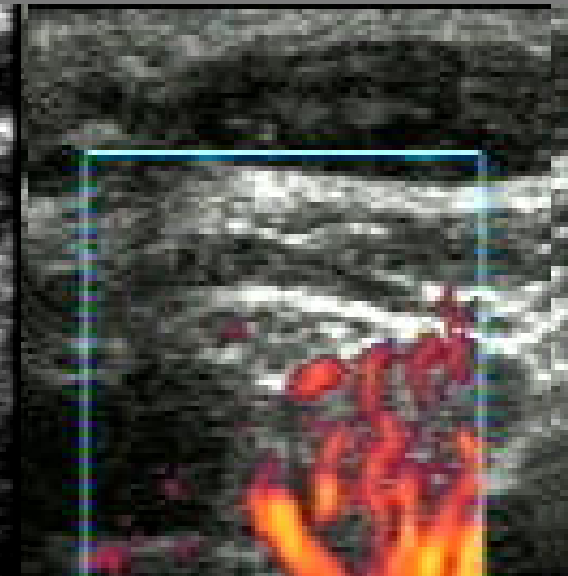
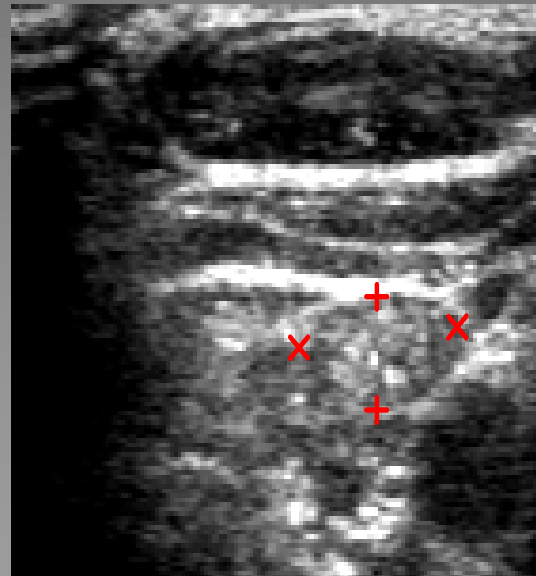
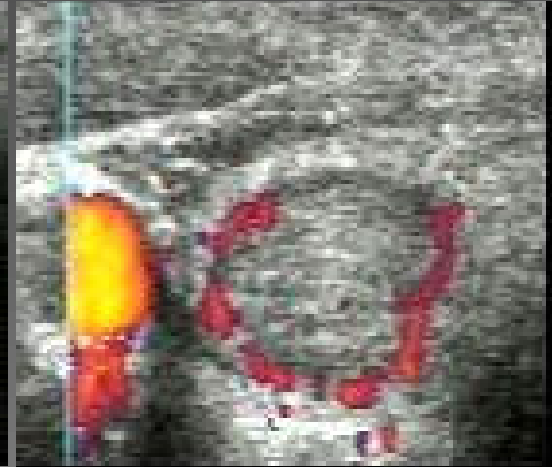
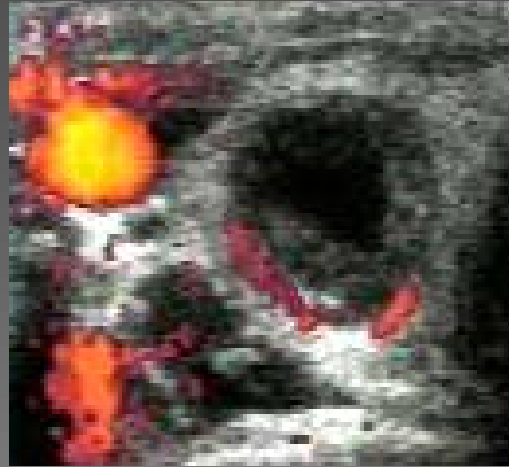
Nodular Goiter / Incidence rate

Statistical data of the Ministry of Public Health

Nodular Goiter in adults - cases per 100 000 persons



Nodular thyroid disease



Incidence of thyroid cancer diagnosed in 1986-2002

Age at exposure (yr)	No. of cases			
	Belarus	Russian Federation	Ukraine	Total
0-14	1,711	349	1,762	3,822
15-17	299	134	582	1,015
Total	2,010	483	2,344	4,837

*UNSCEAR 2008 Report, Annex D:
Over 6,000 thyroid cancers by
2006*

Health Effects of the Chernobyl
Accident and Special Health Care
Programmes / Report of the UN
Chernobyl Forum, 2006

The Belarus-USA Study

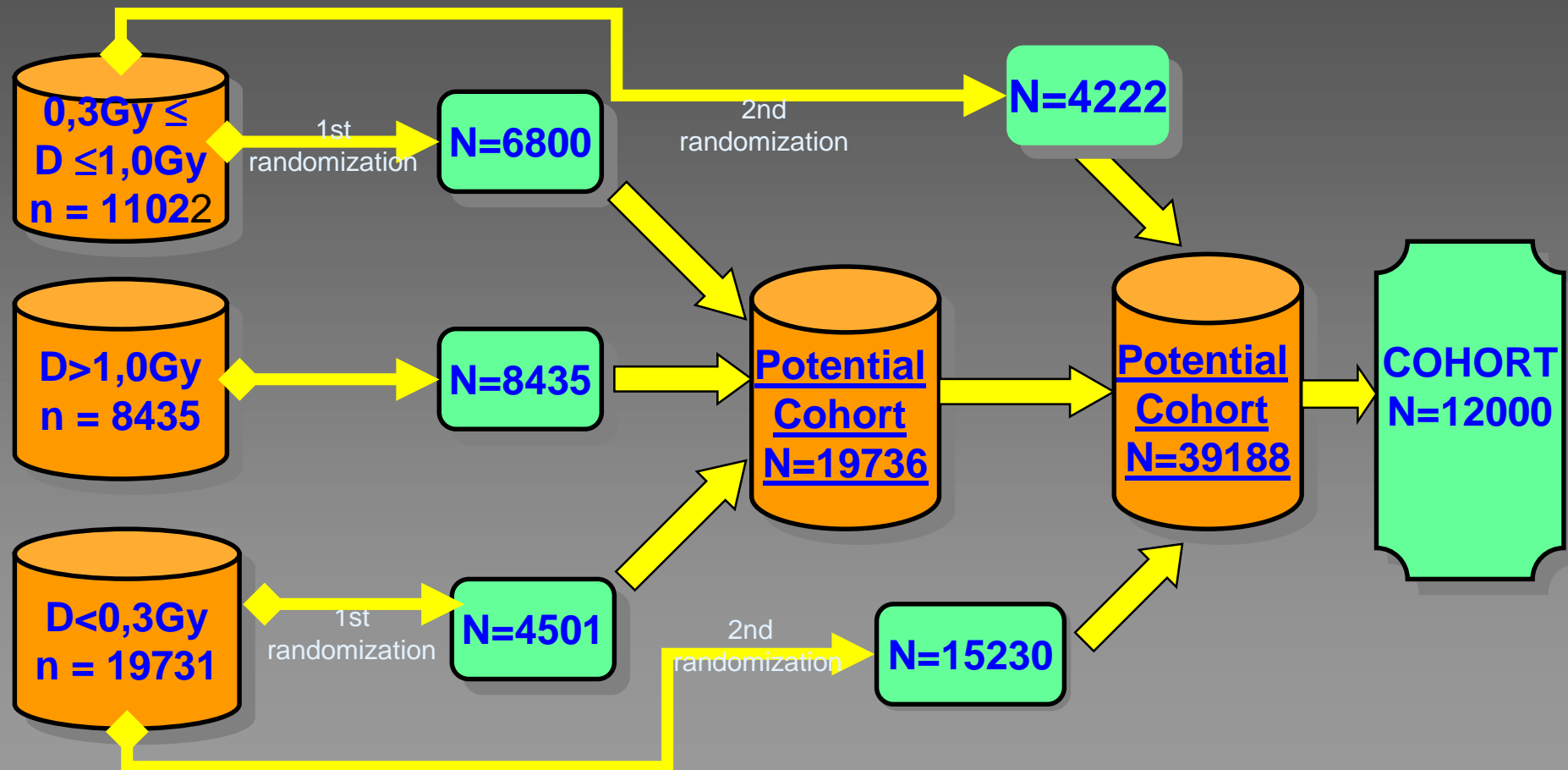
There has been a great increase in the incidence of thyroid cancer and benign thyroid nodules following the release of radioiodine by the Chernobyl accident, which occurred on 26 April 1986. Less thoroughly evaluated has been the prevalence of thyroid autoimmunity, even though external irradiation is recognized to increase the incidence of autoimmune thyroiditis. The Belarus-USA Study was established to quantitate the risk of thyroid and parathyroid disease in a well-defined cohort of individuals under age 18 years at the time of the accident who had direct thyroid radiation measurements. Subjects are examined at least biennially and undergo diagnostic procedures including thyroid palpation and ultrasound (US) and assay of serum for thyroid antibodies, TSH, free T4, and thyroglobulin. By 1 October 2000, 9400 cohort members have been screened at least once.

PREVALENCE OF THYROID ANTIBODIES IN THE BELARUS-USA COHORT STUDY OF THYROID CANCER AND OTHER THYROID DISEASES FOLLOWING THE CHERNOBYL ACCIDENT/ ITC, Kyoto

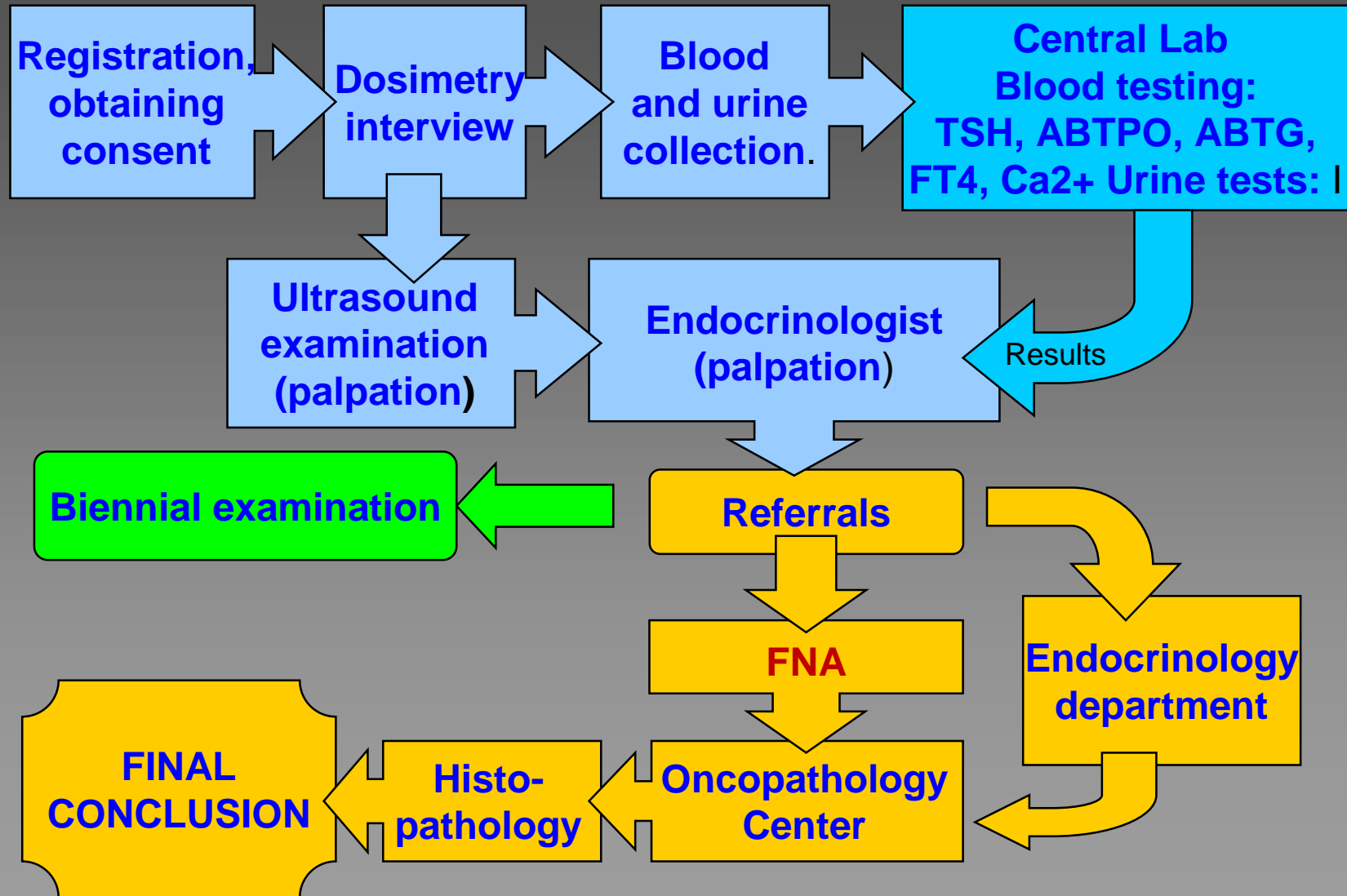
V. Ostapenko, S. Petrenko, O. Polyanskaya, V. Rzhetski, N. Litvinova, N. Lesnikova, E. Buglova, V. Drozd, L. Danilova, V. Stezhko, G.W. Beebe, A.B. Brill, D. Fink, E. Greenebaum, G. Howe, R. McConnell, I. Masnyk, J. Robbins Research Clinical Institute of Radiation Medicine and Endocrinology, Ministry of Health, Belarus; National Cancer Institute, National Institute of Health, Rockville, MD, USA; Columbia University, New York, NY, USA.

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SELECTION OF THE Bel-Am -COHORT



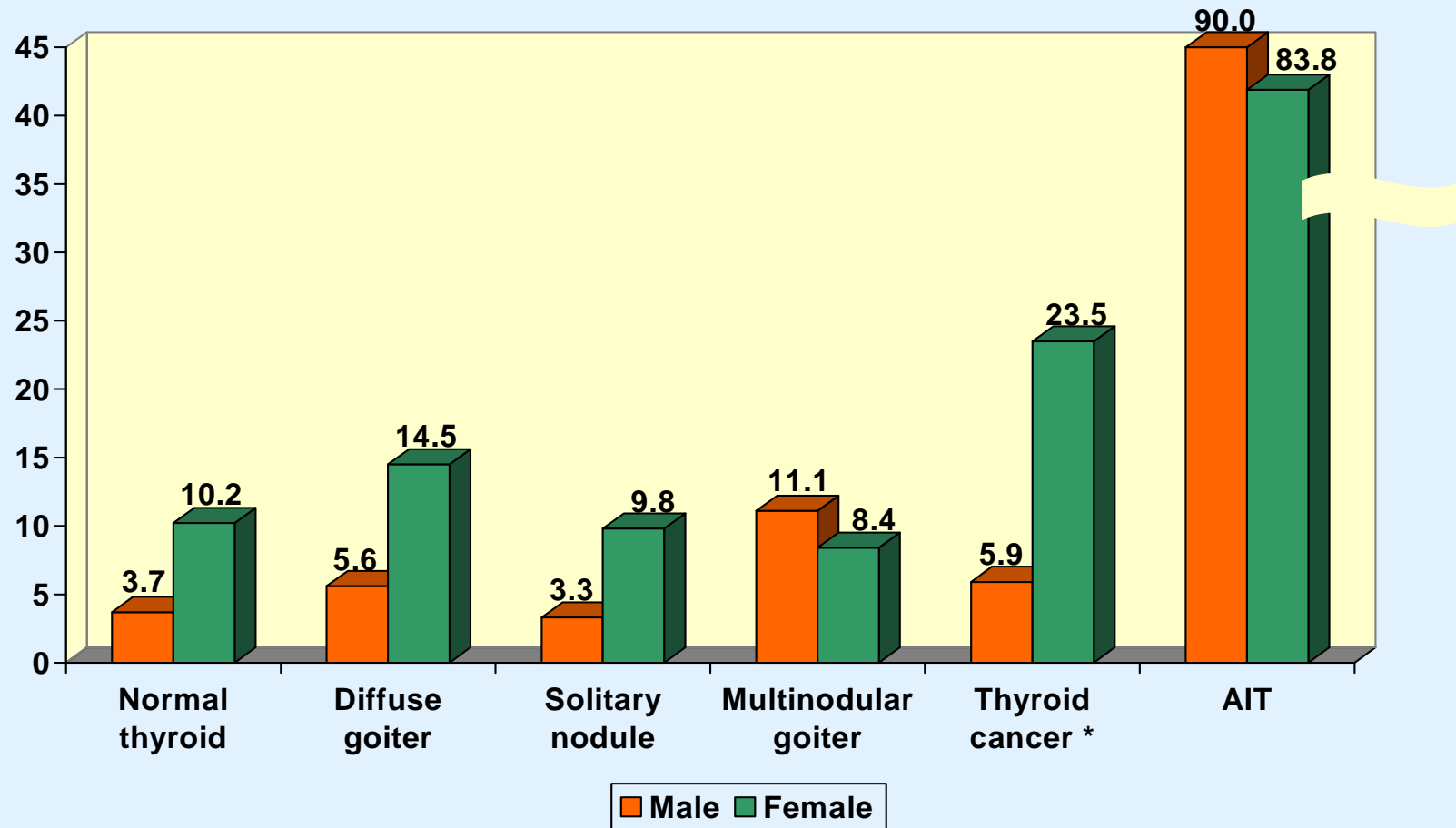
MEDICAL EXAMINATION CHART



PREVALENCE OF ANTITHYROID ANTIBODIES IN EXPOSED SUBJECTS OF BELARUS AND RUSSIA

<i>Characteristics</i>	<i>BelAm</i>	<i>Sasakawa Project</i>	<i>F.Pacini et al., 1998</i>	<i>F.Vermiglio et al., 1999</i>
<i>Age in 1986, years</i>	0 - 18	0 - 10	<1 - 10	In utero - 3
<i>Age at examination, years</i>	11 - 32	5 - 20	6 - 17	5 - 15
<i>Year of examination</i>	1997-2000	1991-1996	1992-1994	1997
<i>Prevalence of ABTPO</i>	5.8%	2.5%	16.7%	18.9%
<i>Prevalence of ABTG</i>	4.9%	1%	8.4%	12.6%

PREVALENCE OF AB(+) IN SUBJECTS WITH THYROID PATHOLOGY



** All diagnoses but thyroid cancer made according to preliminary endocrinological conclusion. Thyroid cancer pathologically confirmed*

Y axis - % AB(+)

Consequences of Chernobyl Accident

All EHP content is accessible to individuals with disabilities. A fully accessible (Section 508-compliant) HTML version of this article is available at <http://dx.doi.org/10.1289/ehp.1205783>.

Research | Children's Health

Measures of Thyroid Function among Belarusian Children and Adolescents Exposed to Iodine-131 from the Accident at the Chernobyl Nuclear Plant

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BACKGROUND: Thyroid dysfunction after exposure to low or moderate doses of radioactive iodine-131 (¹³¹I) at a young age is a public health concern. However, quantitative data are sparse concerning ¹³¹I-related risk of these common diseases.

OBJECTIVE: Our goal was to assess the prevalence of thyroid dysfunction in association with ^{131}I exposure during childhood (≤ 18 years) due to fallout from the Chernobyl accident.

METHODS: We conducted a cross-sectional analysis of hypothyroidism, hyperthyroidism, autoimmune thyroiditis (AIT), serum concentrations of thyroid-stimulating hormone (TSH), and autoantibodies to thyroperoxidase (ATPO) in relation to measurement-based ^{131}I dose estimates in a Belarusian cohort of 10,827 individuals screened for various thyroid diseases.

RESULTS: Mean age at exposure (\pm SD) was 8.2 ± 5.0 years. Mean (median) estimated ^{131}I thyroid dose was 0.54 (0.23) Gy (range, 0.001 – 26.6 Gy). We found significant positive associations of ^{131}I dose with hypothyroidism (mainly subclinical and antibody-negative) and serum TSH concentration. The excess odds ratio per 1 Gy for hypothyroidism was 0.34 (95% CI: $0.15, 0.62$) and varied significantly by age at exposure and at examination, presence of goiter, and urban/rural residency. We found no evidence of positive associations with antibody-positive hypothyroidism, hyperthyroidism, AIT, or elevated ATPO.

CONCLUSIONS: The association between ^{131}I dose and hypothyroidism in the Belarusian cohort is consistent with that previously reported for a Ukrainian cohort and strengthens evidence of the effect of environmental ^{131}I exposure during childhood on hypothyroidism, but not other thyroid outcomes.

KEY WORDS: antithyroid antibodies, autoimmune thyroiditis, Chernobyl, Chornobyl, dose response, hyperthyroidism, hypothyroidism, radioiodine, thyroid gland.

Environ Health Perspect 121:865–871 (2013). <http://dx.doi.org/10.1289/ehp.1205783> [Online 7 May 2013]

In a screening study among 12,000 subjects in Ukraine with doses estimated from individual measurements of thyroid radioactivity, significant associations were found between ^{131}I thyroid dose (mean dose of 0.79 Gy) and prevalence of subclinical hypothyroidism (Ostroumova et al. 2009) and antibodies to thyroperoxidase (ATPO) (Tronko et al. 2006a), but not autoimmune thyroiditis (AIT) (Tronko et al. 2006a) or hyperthyroidism (Hatch et al. 2010).

To extend findings from the Ukrainian cohort, we evaluated functional thyroid outcomes in relation to individual ^{131}I thyroid doses in a comparable cohort of exposed children and adolescents from Belarus who were screened for thyroid cancer and other thyroid diseases. The methods used to estimate thyroid doses and to screen for thyroid diseases in Belarus and Ukraine were similar (Stezhko et al. 2004).

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The most severe accident in the history of nuclear industry occurred on 26 April 1986. evidence of a decline in radiation-related risk of thyroid cancer (Brenner et al. 2011).

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PLOS ONE

Impact of Uncertainties in Exposure Assessment on Estimates of Thyroid Cancer Risk among Ukrainian Children and Adolescents Exposed from the Chernobyl Accident

Mark P. Little^{1*}, Alexander G. Kukush^{2,3}, Sergii V. Maslukh², Sergiy Shklyar^{2,3}, Raymond J. Carroll⁴, Jay H. Lubin⁵, Deukwoo Kwon^{1,6}, Alina V. Brenner¹, Mykola D. Tronko¹, Kiyohiko Mabuchi¹, Tetiana I. Bogdanova⁷, Maureen Hatch¹, Lydia B. Zablotska⁸, Valeri P. Tereshchenko⁹, Evgenia Ostroumova¹, André C. Bouville¹, Vladimir Drozdovitch¹, Mykola I. Cherpumy², Lina N. Vokogan², Steven L. Simon¹⁰, Victor M. Shpak², Ilva A. Liktarev²

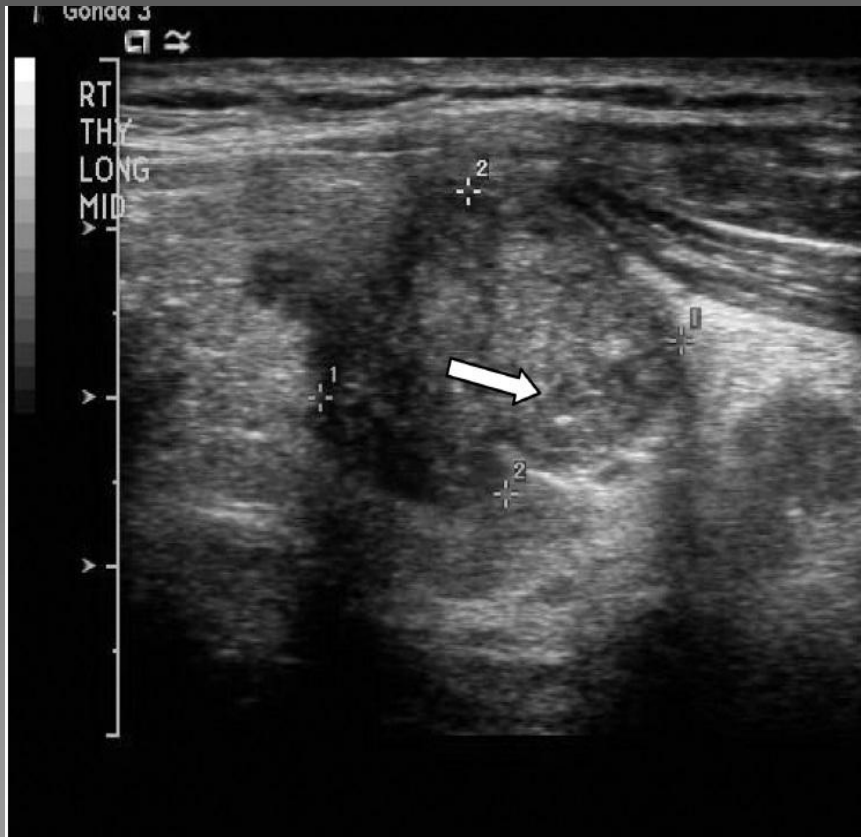
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Abstract

The 1986 accident at the Chernobyl nuclear power plant remains the most serious nuclear accident in history, and some thyroid cancers, particularly thyroid carcinomas, have been exposed to release of radioactive iodine via the air-dispersed regular. Failure to take dose-response information into account can lead to bias in assessments of dose-response relationships. Although risks in the Ukrainian-US thyroid screening study have been previously evaluated, errors in dose assessments have not been addressed hitherto. Dose-response patterns were examined in a thyroid screening prevalence cohort of 43,127 persons aged <16 at the time of the accident, who were resident in the most radioactively contaminated regions of Ukraine. We obtained cohort data on thyroid cancer incidence, thyroid cancer mortality, and thyroid cancer prevalence. We used the maximum-likelihood correction, various types of regression calibration, and Monte Carlo maximum-likelihood, were applied to the doses that can be derived from the ratios of thyroid activity to thyroid mass. The two components that make up this ratio have different types of error. Berkson error for thyroid mass and classical error for thyroid activity. The first regression calibration method yielded estimates of excess odds of 2.78 (95% CI 1.92-47.46), whereas the second method yielded estimates of 4.93 (95% CI 1.67-19.93), both less than unadjusted analysis. The Monte Carlo maximum-likelihood method produced an excess odds ratio of 4.93 (95% CI 1.67-19.93), both less than unadjusted analysis. There are borderline significant ($p=0.101$) 12% indications of downward curvature in the dose response, allowing for which nearly doubled the low-dose hazard coefficients. The results of this study suggest that the use of thyroid activity to thyroid mass ratios in the presence of error in the numerator and errors of exposure of Berkson and classical type, associated with thyroid dose assessments.



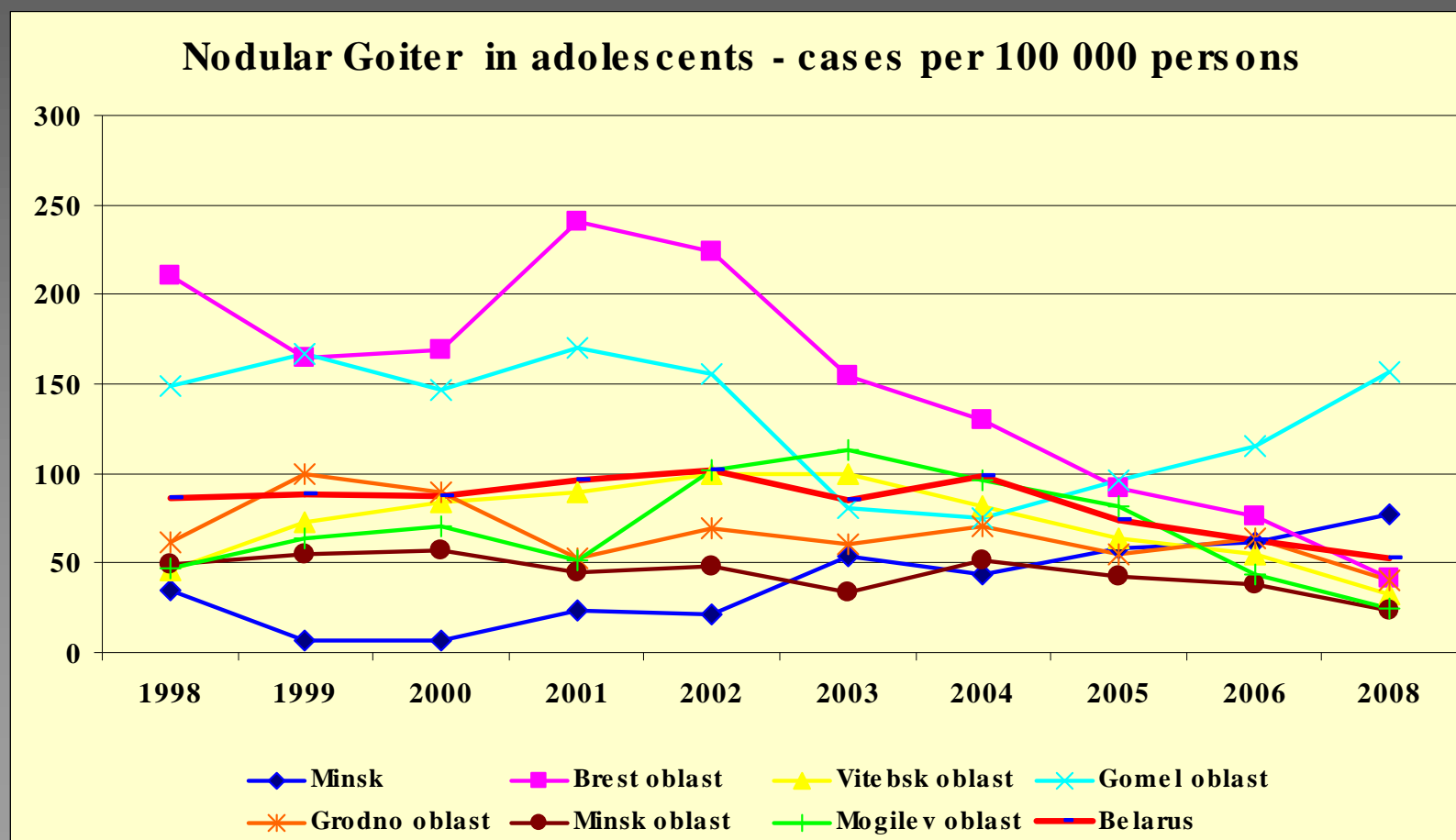
Thyroid cancer



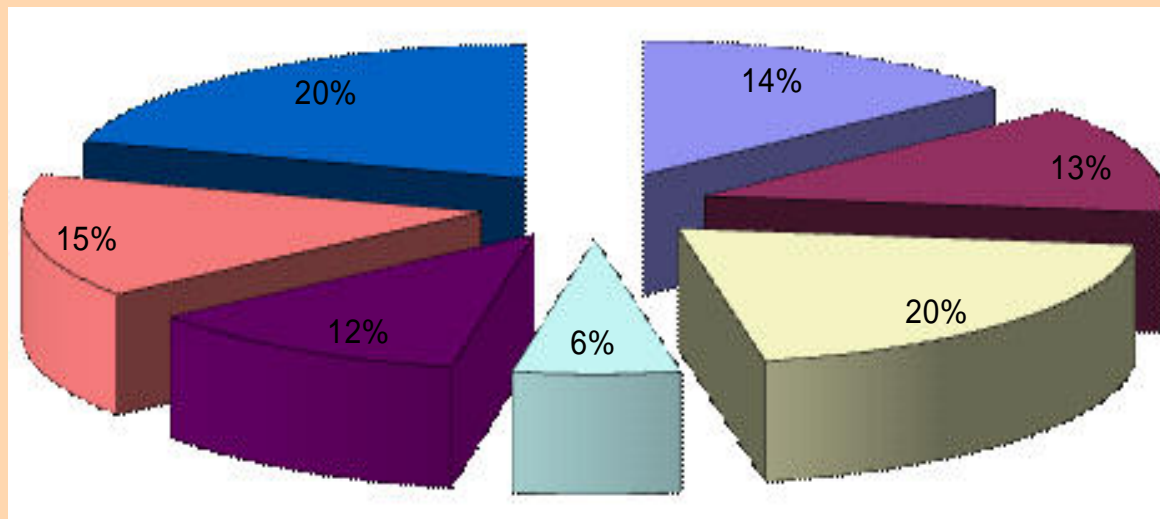
- *Current Belarusian guidelines for the treatment of progressive DTC include optimization of TSH suppression, ¹³¹I, resection of selected metastases, bisphosphonates for bony metastases (Y. Demidchik, L.Danilova, V.Drozd, M.Lushchyk, 2010);*
- *Like the RET/PTC, mutations in BRAF are thought to occur early in thyroid cancers tumorigenesis (Nikiforova et al. 2003) and discussed as an attractive target for molecular therapies.*

Nodular Goiter / Incidence adolescents

Statistical data of the Ministry of Public Health



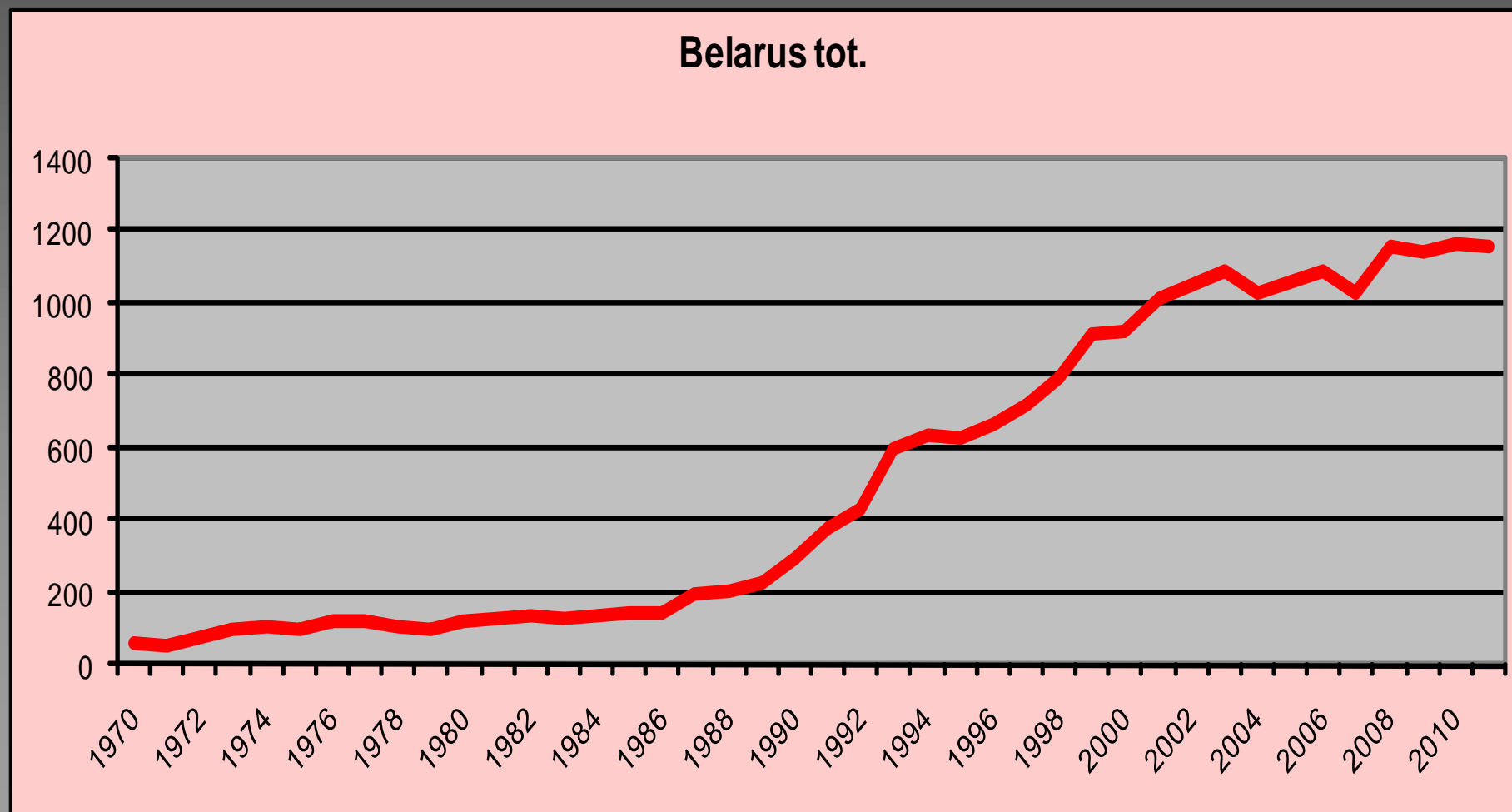
Distribution of total amount of patients with Thyroid Cancer in different regions of Belarus 1990-2011 (21 616 cases)



- Brest rgn.
- Vitebsk rgn.
- Gomel rgn.
- Grodno rgn.
- Minsk rgn.
- Mogilev rgn.
- Minsk city

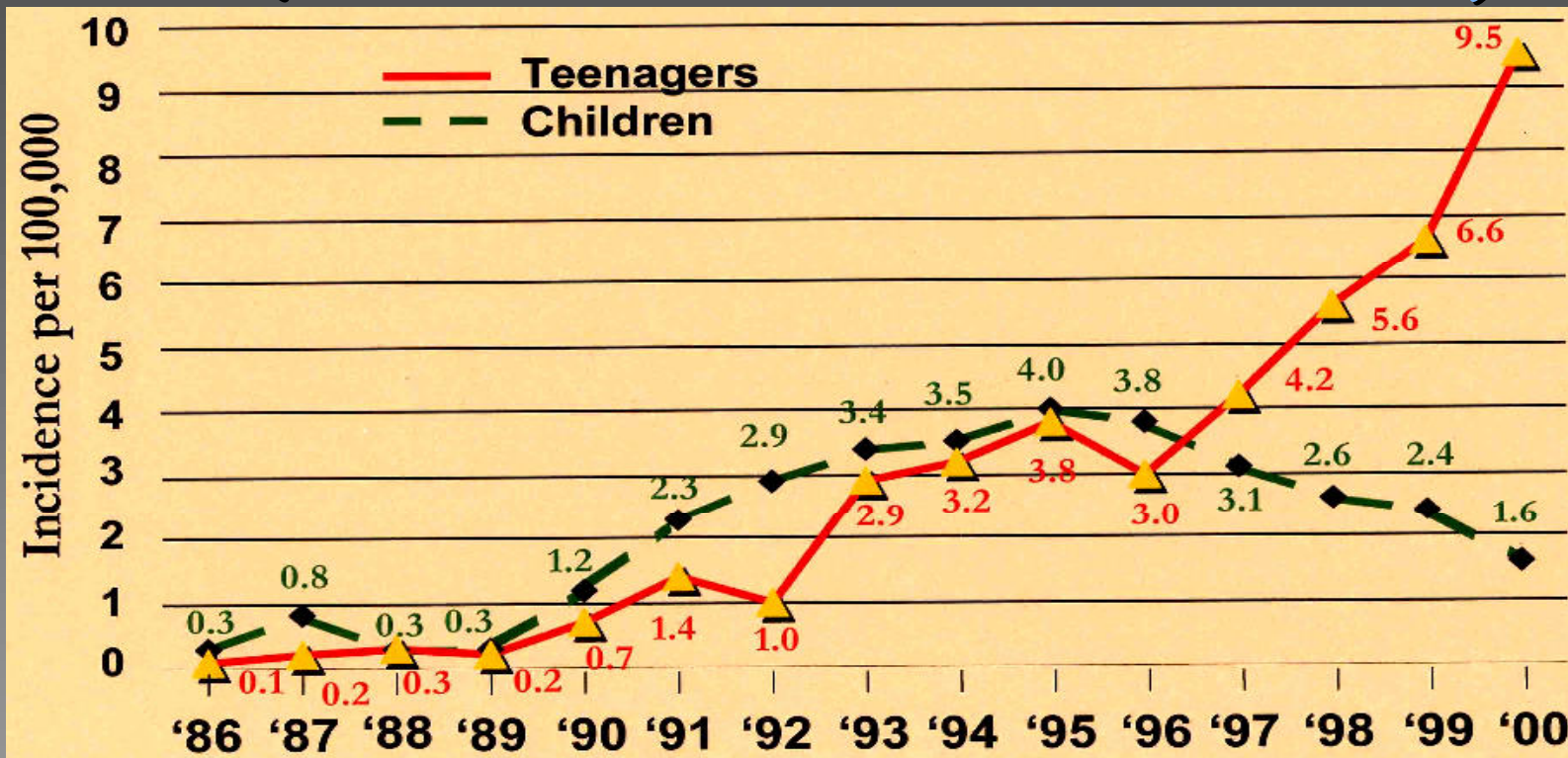
Yearly number of patients with thyroid cancer in Belarus: 1990 to 2011

Statistical data of the Ministry of Public Health



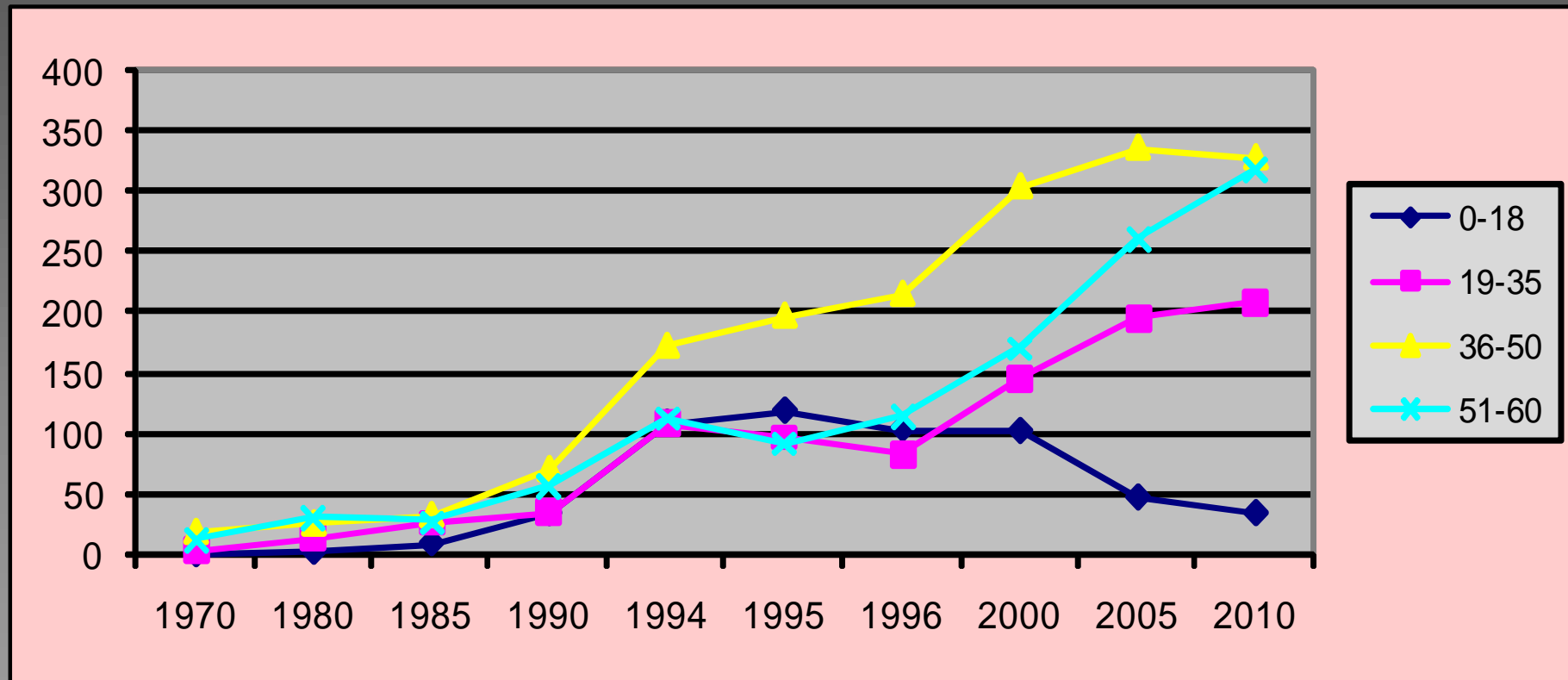
Thyroid Cancer in Children and Teenagers

(E.P. Demidchik et al, 2002-2004)

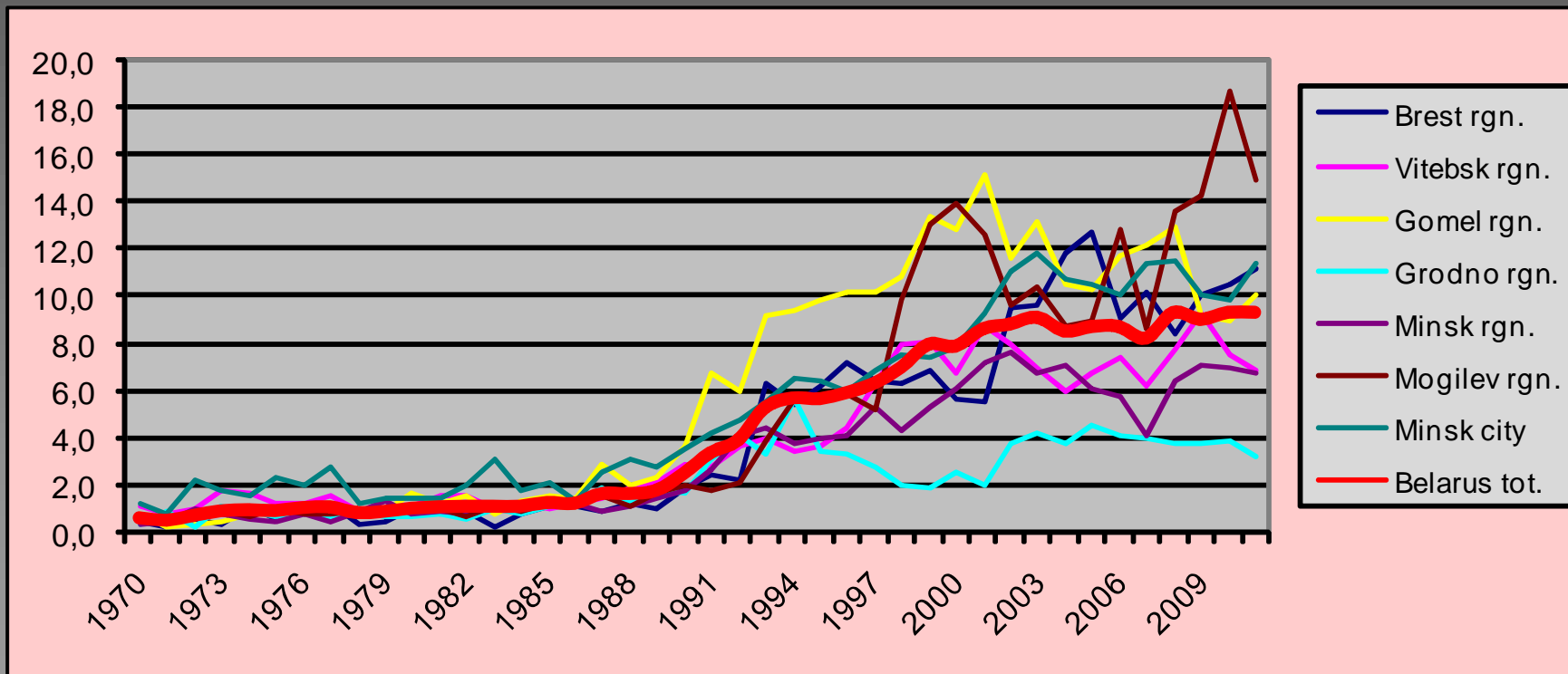


Thyroid cancer: Age-dependent incidence

(Y. Demidchik et al, 2012)

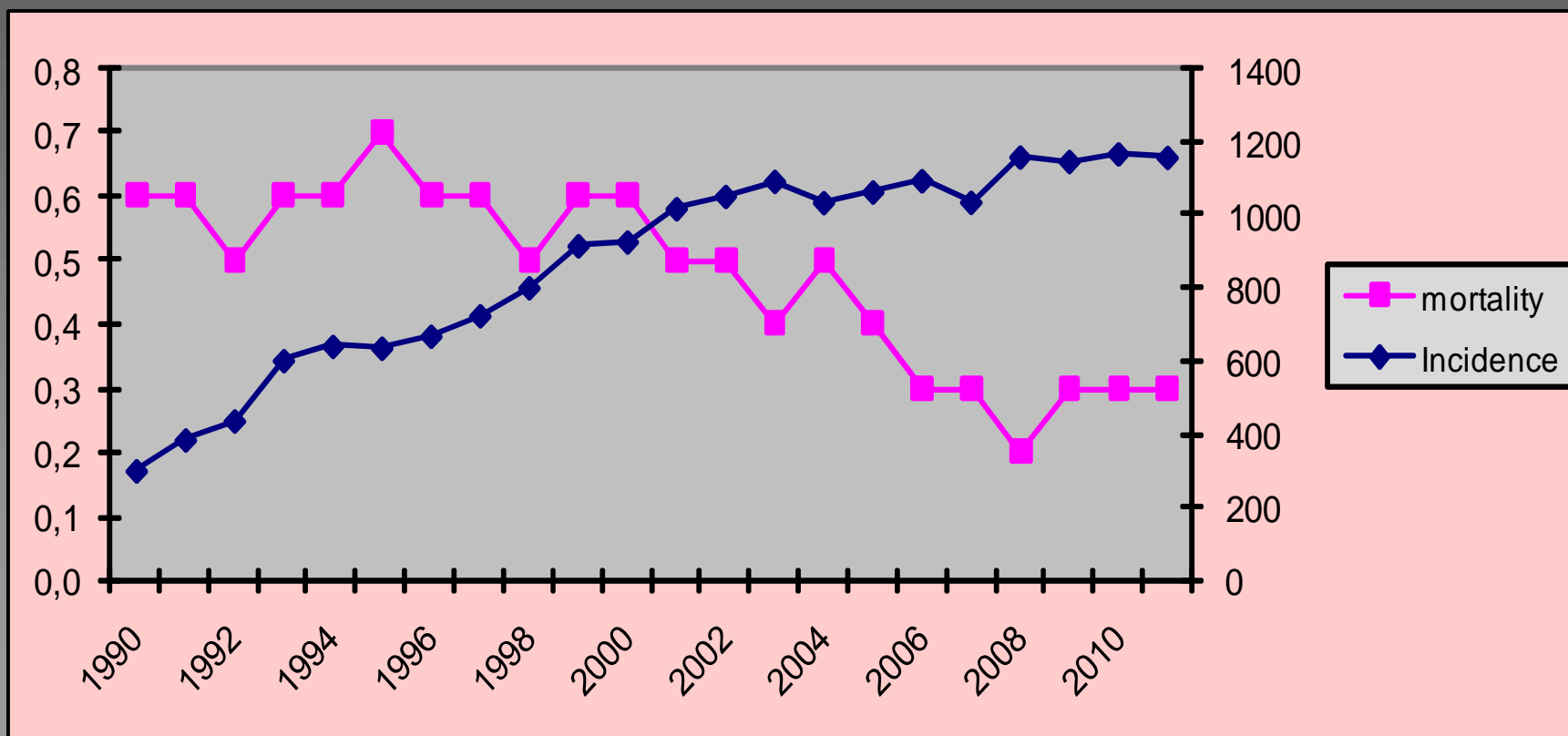


Statistical data of the Ministry of Public Health

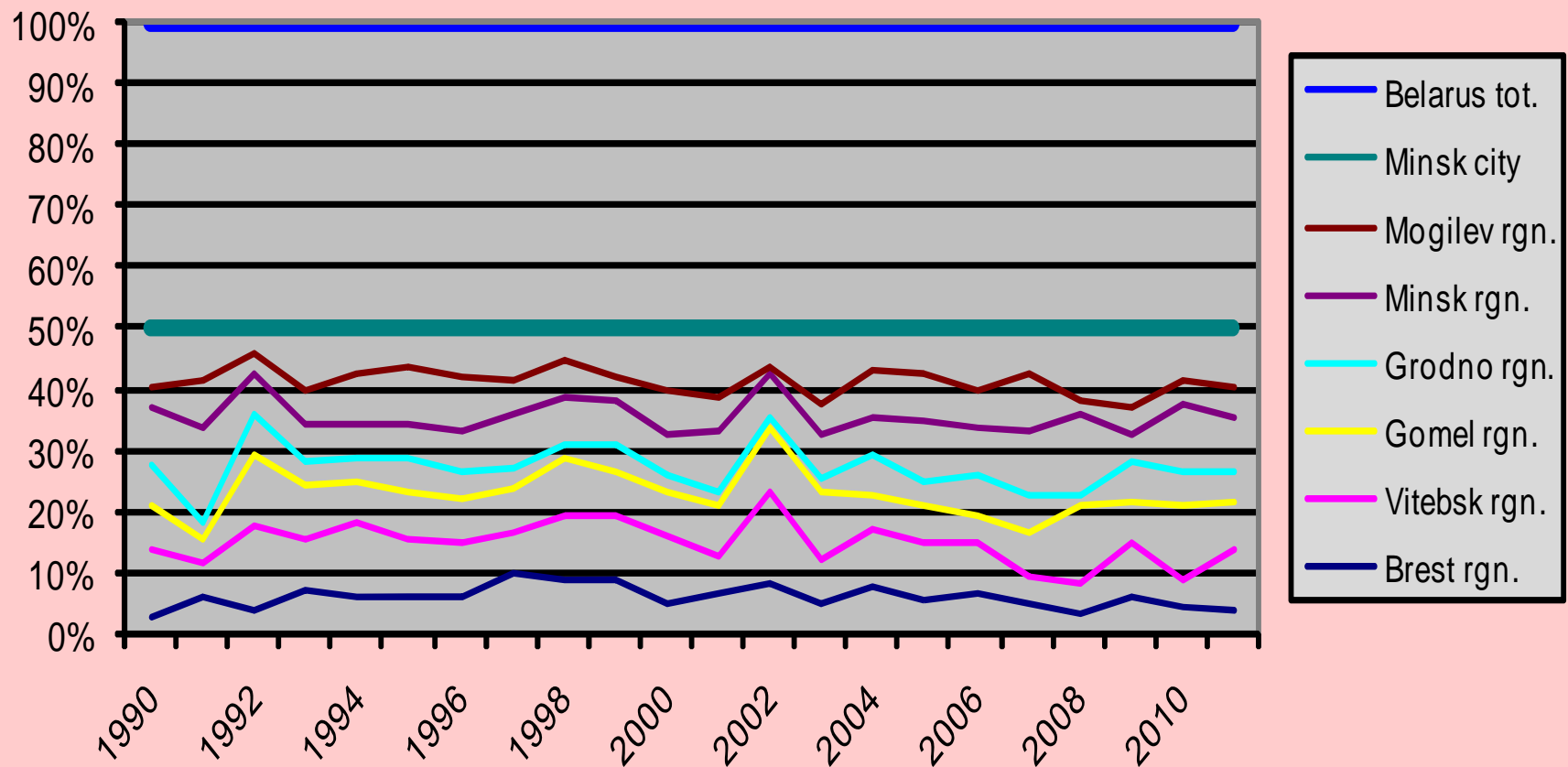


Thyroid cancer : Mortality and Incidence rate

Statistical data of the Ministry of Public Health

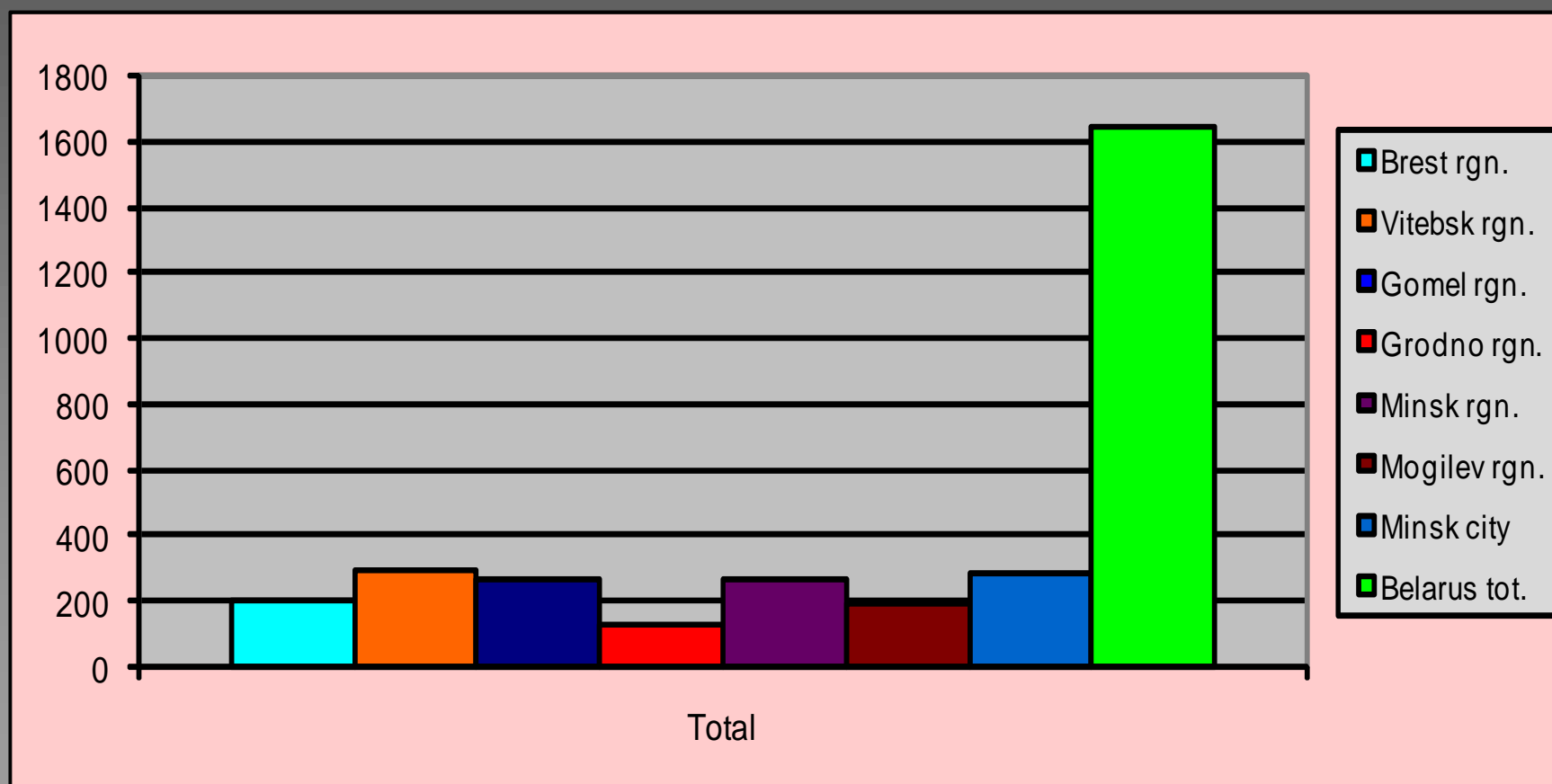


Thyroid cancer : mortality since 1990, absolute number of cases (normalized graph) *(Y. Demidchik et al, 2012)*



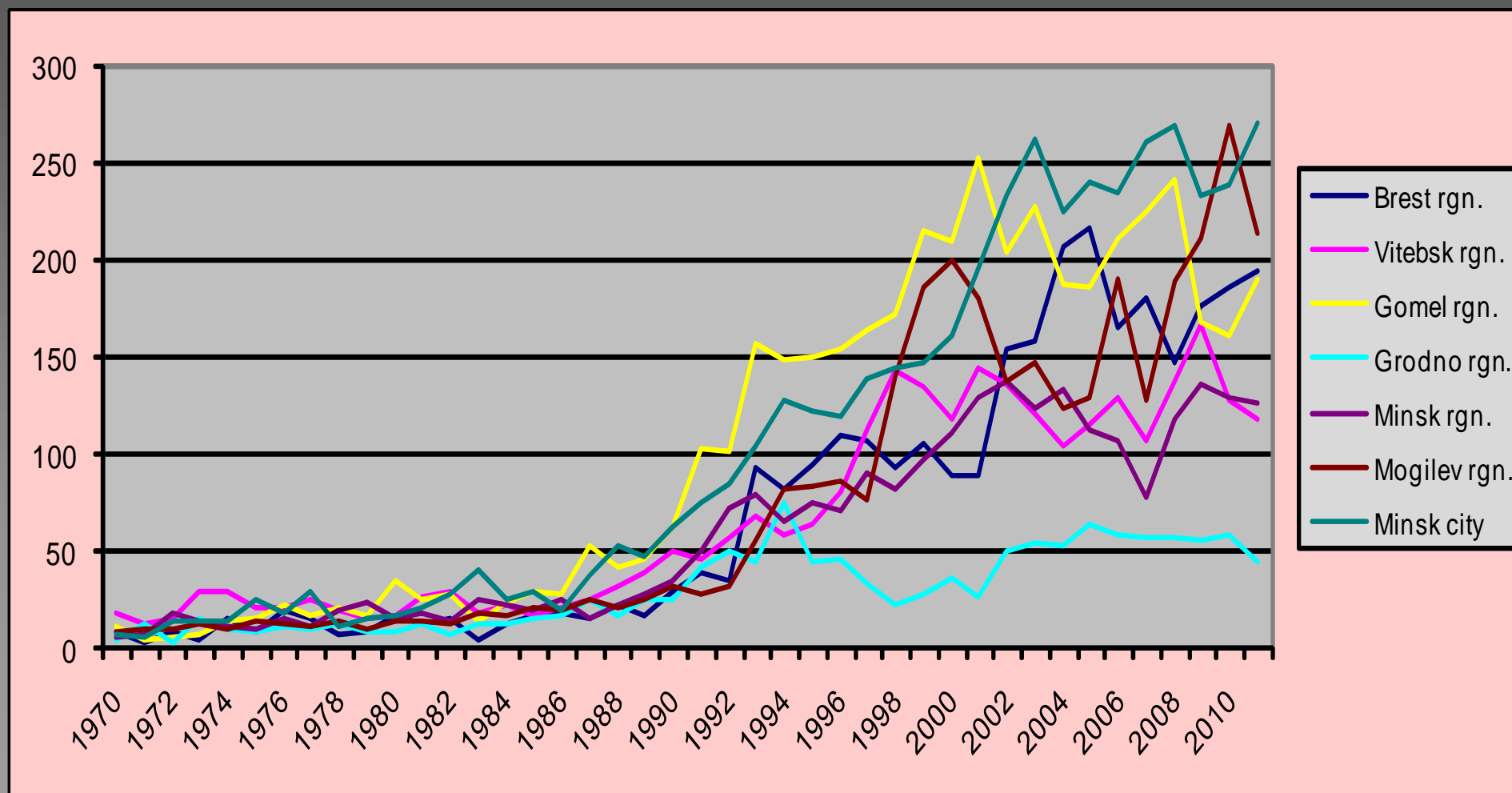
Thyroid cancer: mortality, total number of deceased 1990-2013

Statistical data of the Ministry of Public Health



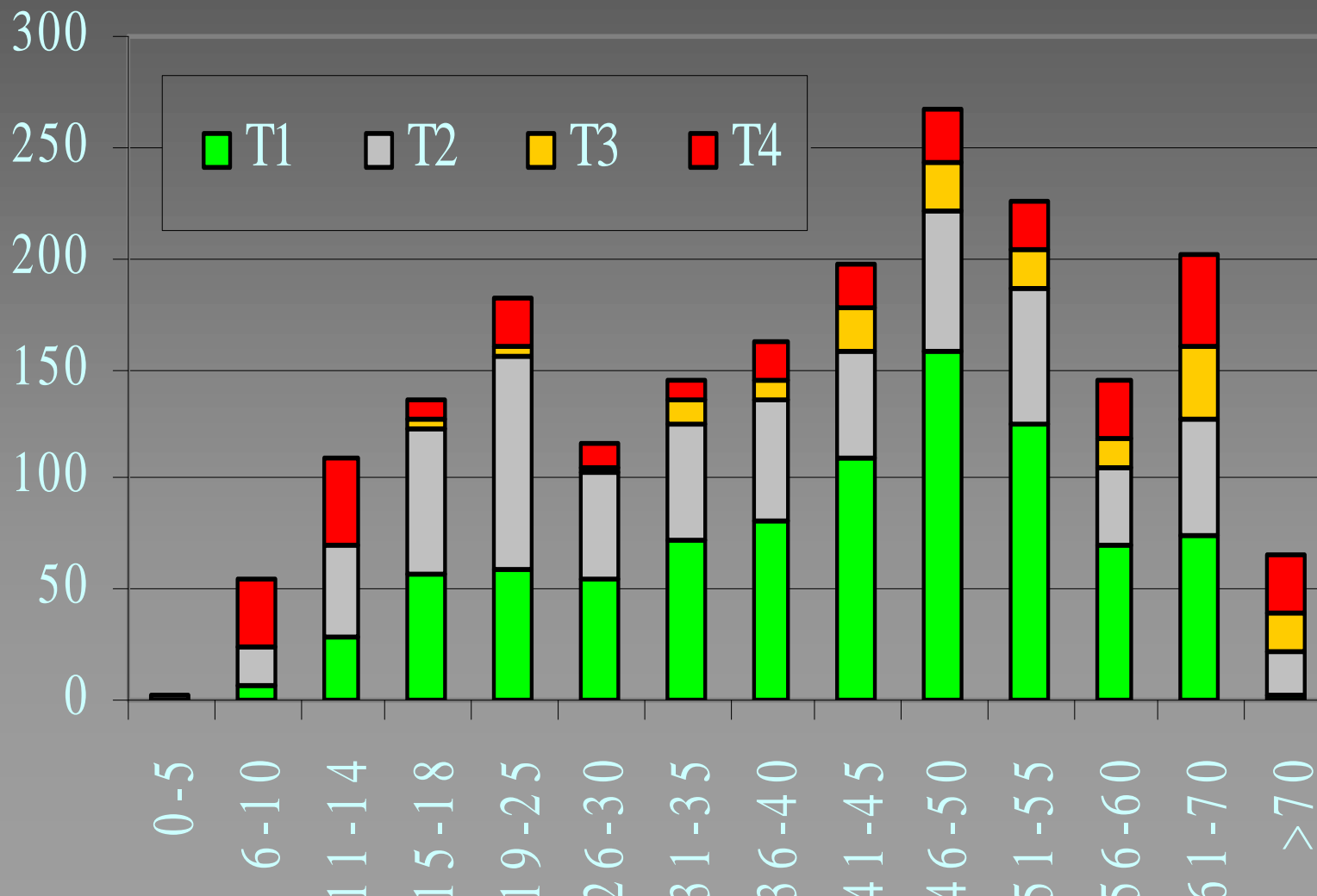
Yearly number of patients with thyroid cancer 1990-2011 in Belarus

Statistical data of the Ministry of Public Health



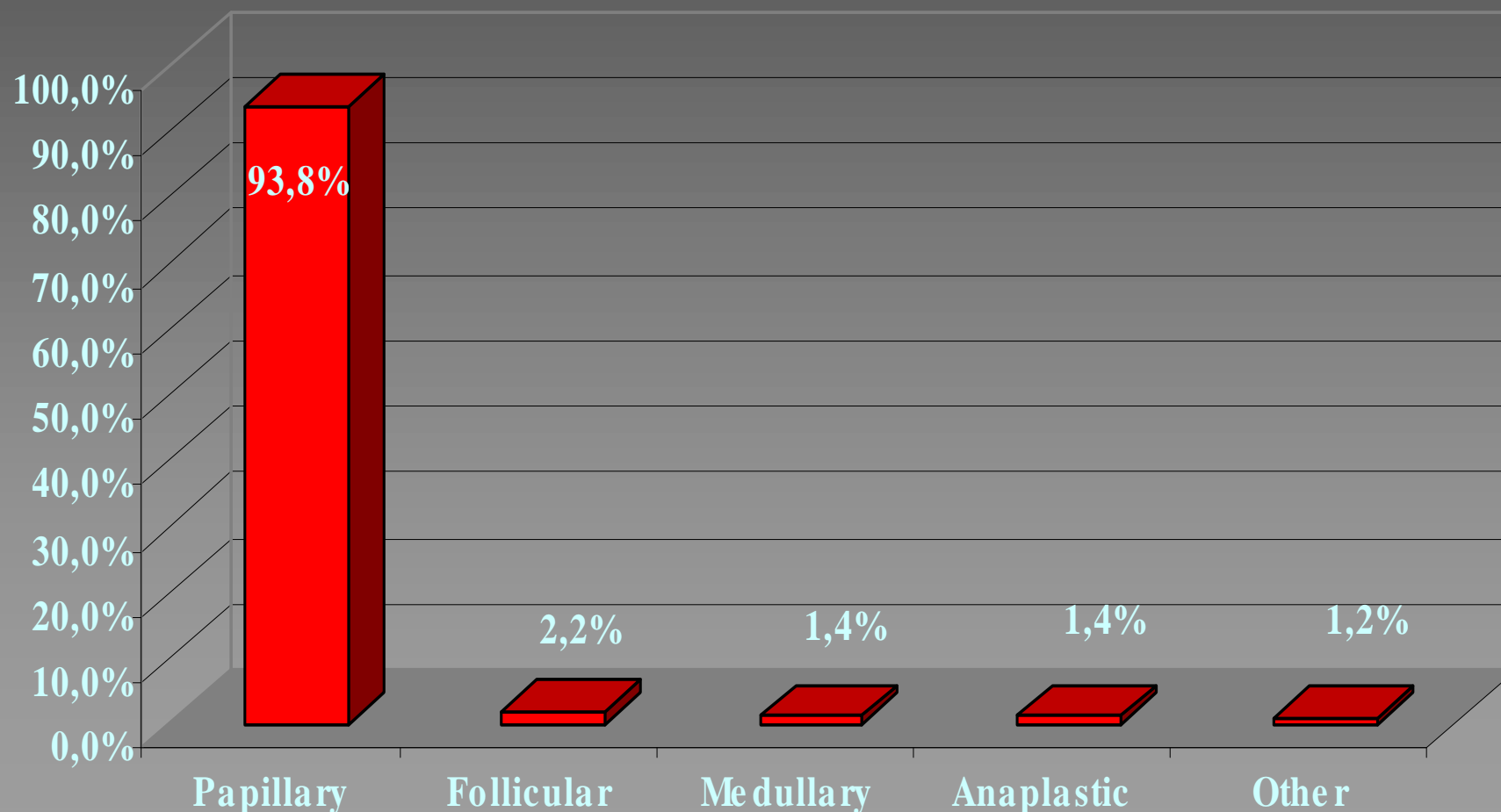
Distribution of Thyroid Cancer cases in different age groups according to TNM stages

Brest region n=2005 / Brest Regional Dispensary statistic data



Percentage ratio of different morphological variants of thyroid cancer in Brest region 1990-2013

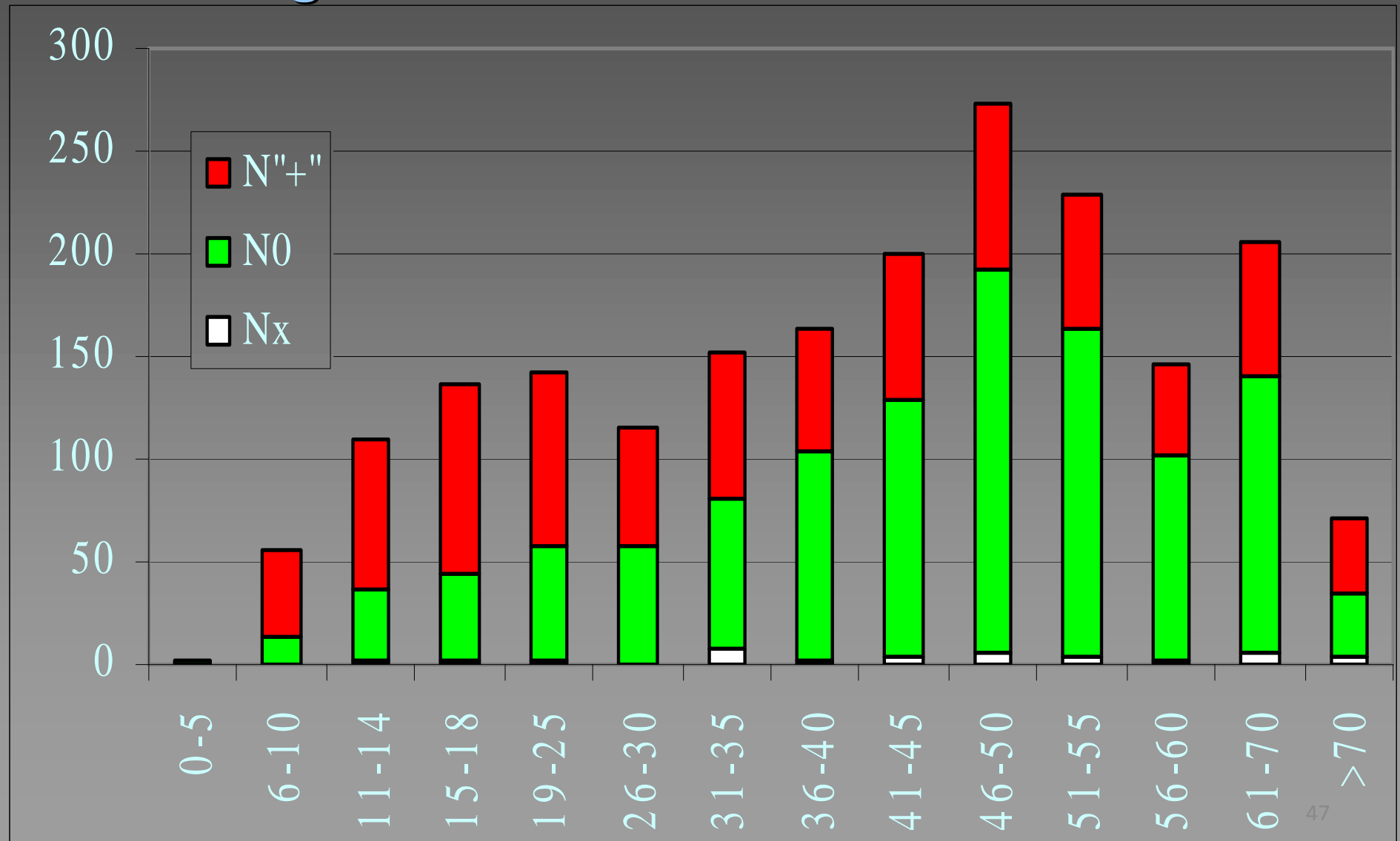
Brest Regional Dispensary statistic data



Distribution of Thyroid Cancer cases in different age groups according to TNM stages

Brest region

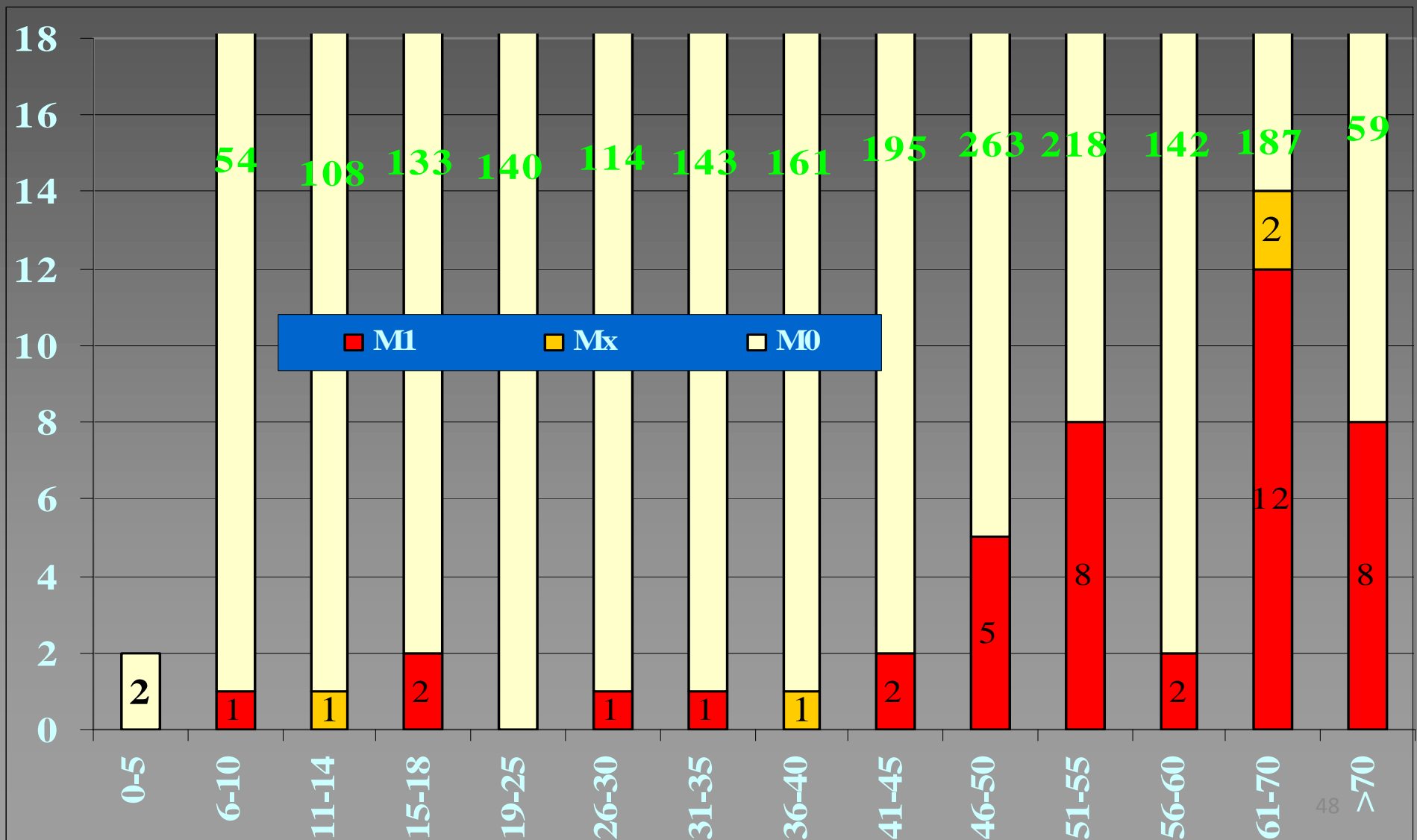
n=2000



Distribution of Thyroid Cancer cases in different age groups according to TNM stages

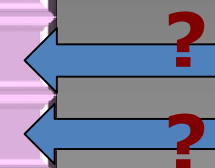
In Brest region

n=1965



Thyroid average and collective doses /¹³¹Iodine and number of thyroid cancer cases in Republic of Belarus

Contaminated territory	Average dose (mGy)			Collective dose	Number of Thyroid Cancer cases	
	0-7 years	14-18 years	adults		Predicted	Real to 2005
Minsk oblast	22,9	7,1	7,4	14 530	101	124
Gomel oblast	475,8	145,0	148,1	321 750	569	509
Mogilev oblast	97,6	29,4	30,7	50 020	173	55
Brest oblast	77,8	23,9	24,7	45 170	49	223
Grodno oblast	16,7	5,2	5,4	7 780	20	49
Vitebsk oblast	5,5	1,6	1,7	2 720	8	16
Belarus	122	37	37	476 000	920	976



Gavrilin, Y., V. Khrouch, S. Shinkarev et al. Individual thyroid dose estimation for a case-control study of Chernobyl-related thyroid cancer among children of Belarus-part I: ¹³¹I, short-lived radioiodines (¹³²I, ¹³³I, ¹³⁵I), and short-lived radiotelluriums (^{131m}Te and ¹³²Te). Health Phys. 86(6): 565-585 (2004).

Gavrilin, Y.I., V.T. Khrouch, S.M. Shinkarev et al. Chernobyl accident: reconstruction of thyroid dose for inhabitants of the Republic of Belarus. Health Phys. 76(2): 105-119 (1999).

Thyroid cancer in Belarus

- The incidence of thyroid cancer in children is likely to be decreased;
 - 2007- 11 newly diagnosed cases,
 - 2010 – 11 cases,
 - 2011 - 19 cases;
- *The incidence of thyroid cancer among older age groups of population is increasing*
 - 1993 – 484 cases
 - 2010 – 1098 cases
 - 2011 - 1165

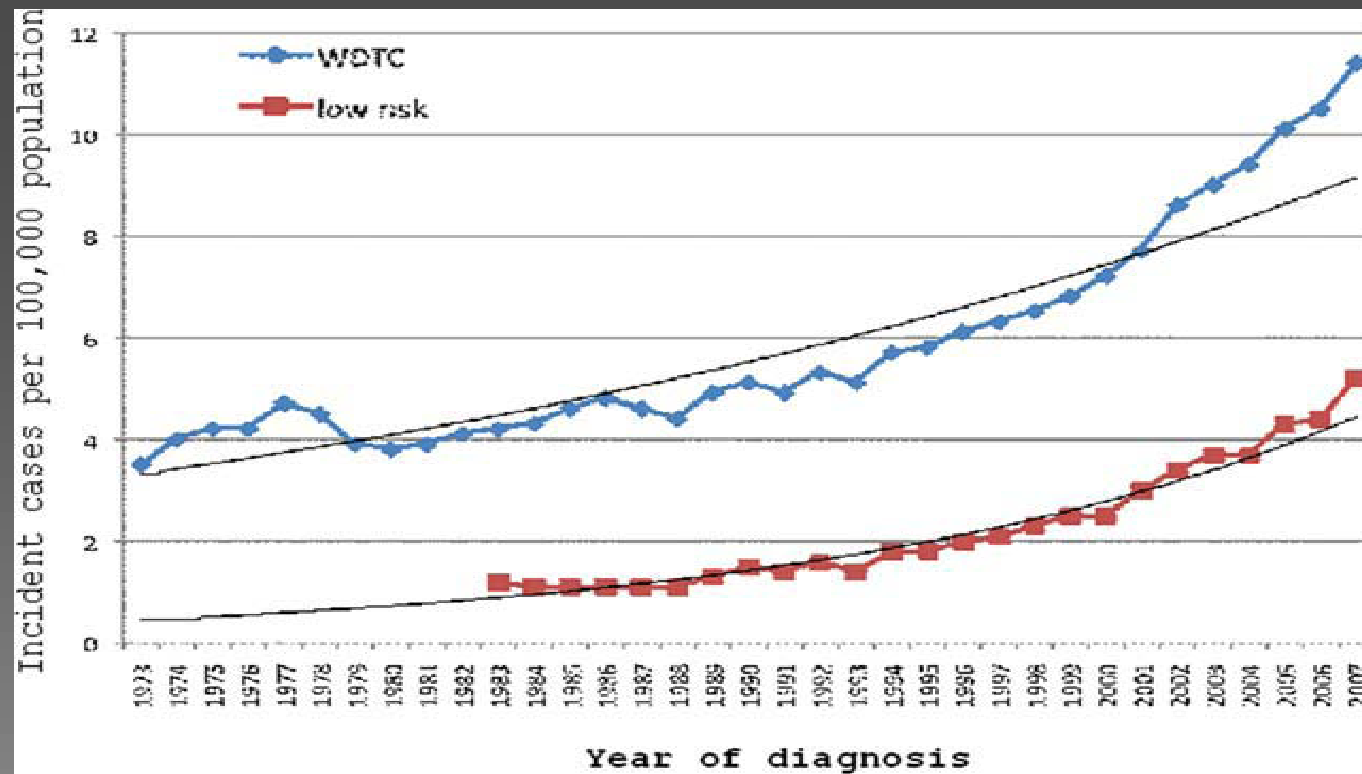
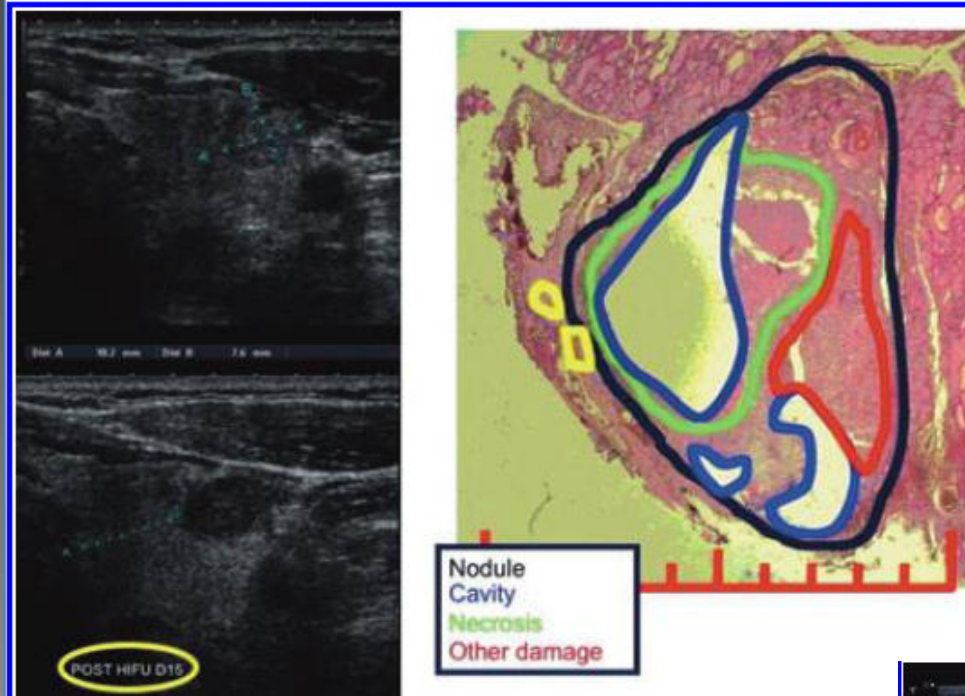


Fig. The incidence of well differentiated thyroid cancer (WDTC) over time is illustrated for all patients and for low risk patients

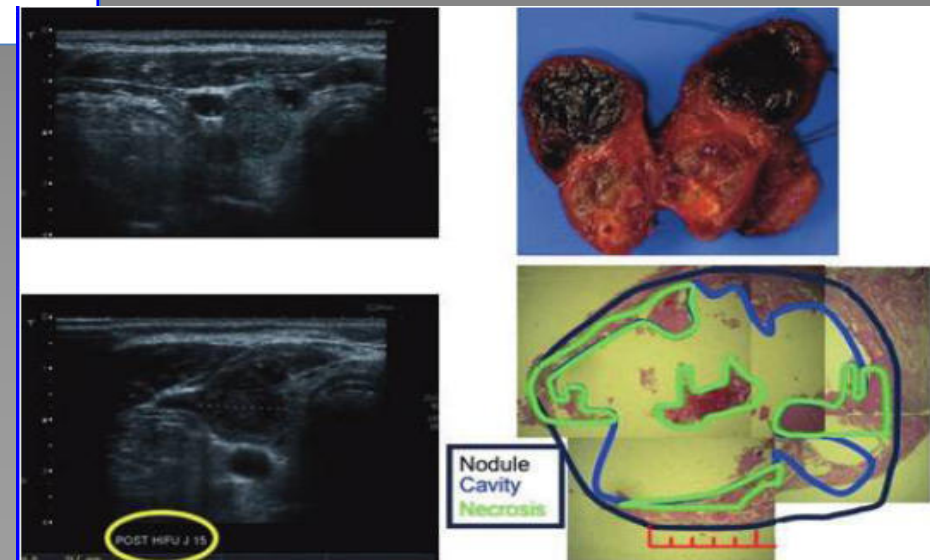
Gopalakrishna Iyer N, Morris L, Michael Tuttle R, ; Shaha A, Ganly I, Rising Incidence of Second Cancers in Patients With Low-Risk (T1N0) Thyroid Cancer Who Receive Radioactive Iodine Therapy. Cancer 2011; Published online in Wiley Online Library (wileyonlinelibrary.com)

HIFU TREATMENT OF THYROID NODULES



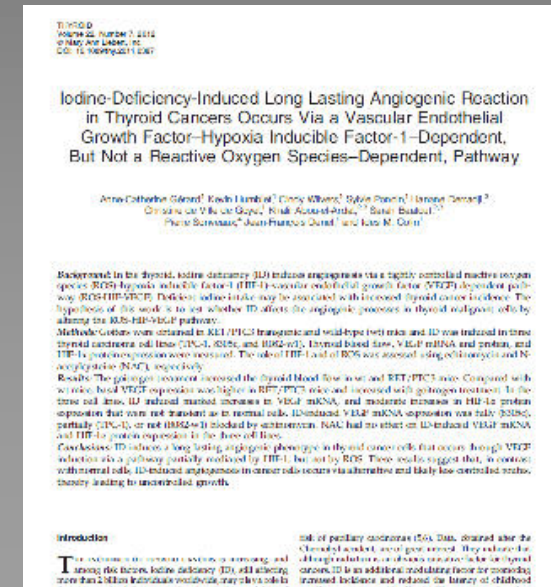
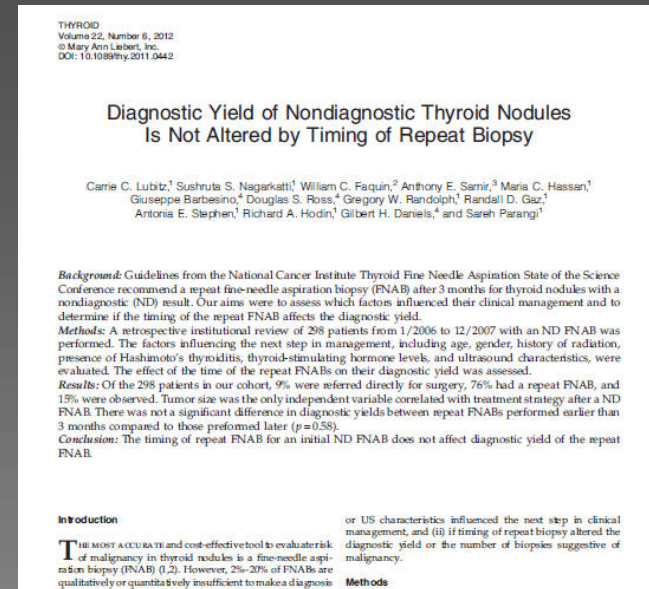
- New ablation technologies in Nodular Thyroid diseases

Conservative Treatment of Thyroid Nodular Disease



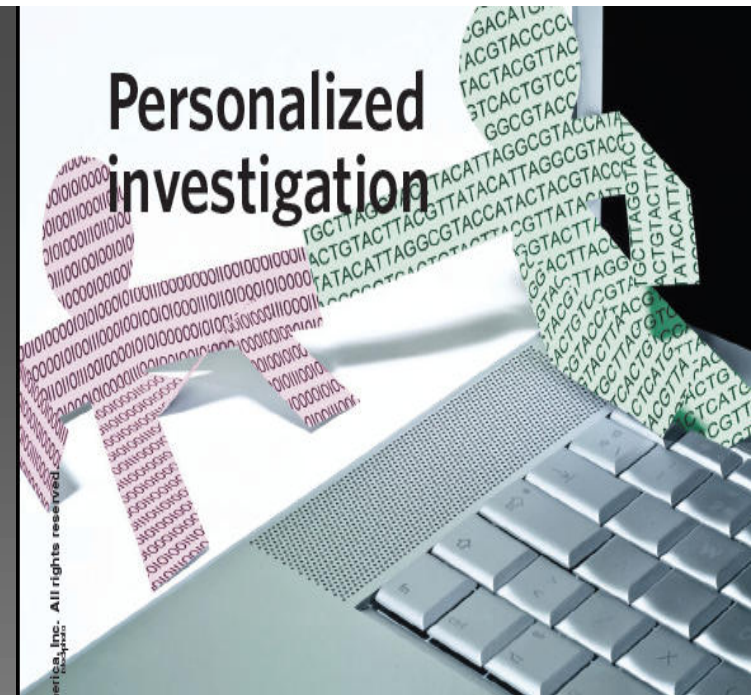
Thyroid nodules

- ***BRAF was found to be mutated (BRAFFV600E) in 28–69% of PTC (Cohen et al. 2003, Fukushima et al. 2003, Kimura et al. 2003, Namba et al. 2003, Nikiforova et al. 2003, Soares et al. 2003, Xu et al. 2003, Trovisco et al. 2004, Xing et al. 2004).***
- ***BRAF is a non-receptor serine threonine kinase involved in the RAS/RAF/MAPK/ERK signaling cascade***



Personalized Thyroidology

- **Personalized prophylaxes**
- **Personalized diagnosis**
- **Personalized treatment**
- **Personalized prognosis**



Despite continued doubts about the clinical utility of direct-to-consumer genetic tests, tens of thousands of people have sent away tubes full of their saliva to learn more about their genetic profiles. Armed with such DNA data, a number of early adopters are showing how empowering—and beneficial to science—personal genetic information can be. Elie Dolgin reports on one company's plans to make medical genetics more participatory.

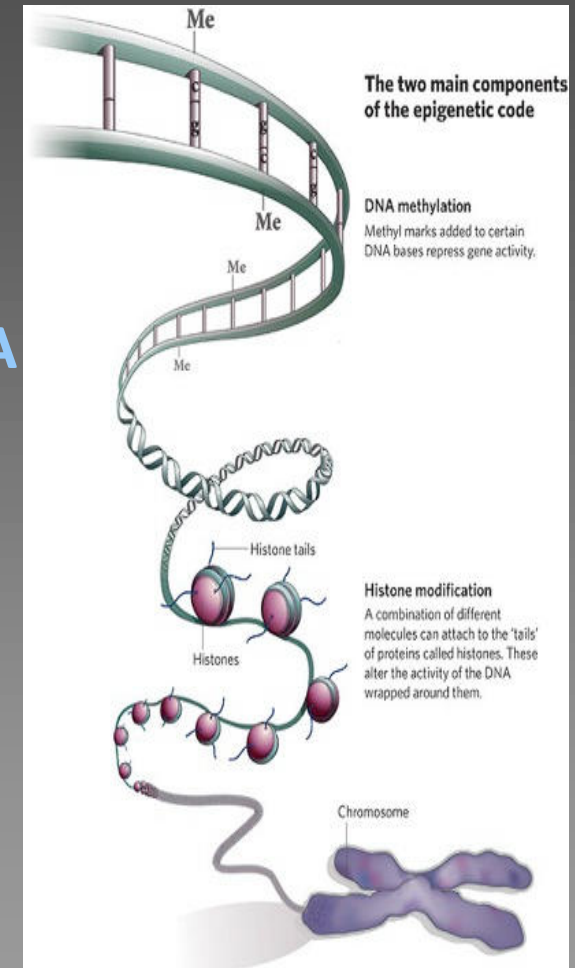
Three years ago, the DNA technology provider Illumina teamed up with the company 23andMe to develop genotyping chips for the latter's direct-to-consumer gene test kits. To celebrate the partnership, Illumina employees had the rare opportunity to order discounted gene tests for just \$249—a low cost compared to the \$999 price tag at the time.

Even after shedding close to a third of his body weight, McCauley still wanted to do more with the gene profile data—a collection of around 600,000 single nucleotide polymorphisms (SNPs) associated with various ancestries and disease risks. Yet, beyond the obvious lifestyle

Now, with an explosion of companies offering personalized genetic tests, the intersection between genetics and medicine is starting to be tapped by amateur biologists-at-large as

Epigenetic therapy and prophylaxes

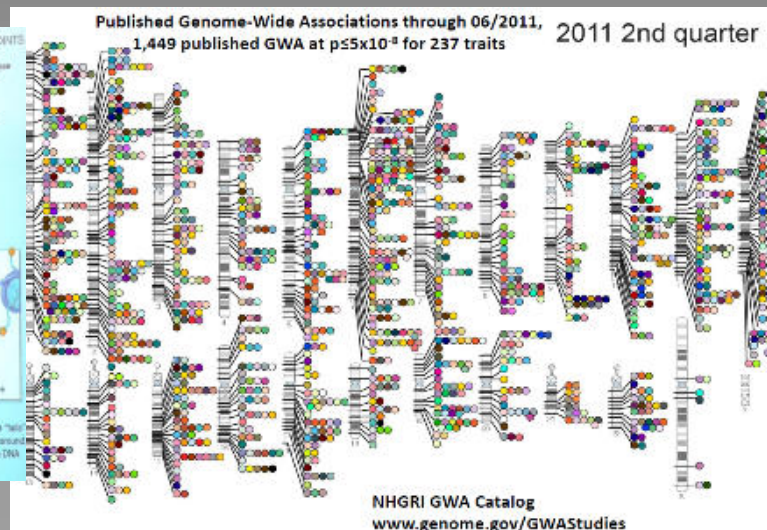
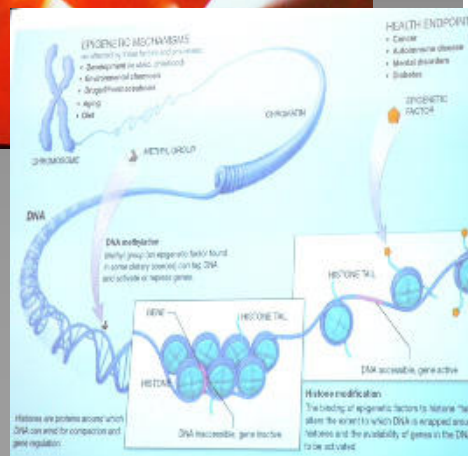
- The complexity of human carcinogenesis cannot be accounted for by genetic alterations alone, but also involves epigenetic changes in processes such as DNA methylation, histone modifications, and microRNA expression.
- In turn, these molecular alterations lead to permanent changes in the expression of genes that regulate the neoplastic phenotype, such as cellular growth and invasiveness.
- *Targeting epigenetic modifiers has been referred to as epigenetic therapy*



R.Taby, J.-P. J. Issa, Cancer J Clin 2010; W.G. Feero, et al. NEJM 362; 2010

Thyroid cancer in Belarus: Personalized prognosis, diagnosis and treatment

- *Environmental chemicals*
- *Drugs/Pharmaceuticals*
- *Combined effect of radiation exposure and environmental factors*
- *Development in utero and in childhood*



Conclusion

- *Continuation of active screening programs in contaminated and non contaminated regions of Belarus are required to elucidate the prevalence of widespread thyroid dysfunctions, early detection of thyroid cancer in high risk groups (those who had radiation exposure to the thyroid gland in utero and childhood, people living in the areas around Chernobyl or moved to other places after they had received radiation doses), and verified influence of endocrine disrupters in modifying epigenetic mechanisms or potentiate the carcinogenic effect of radiation exposure*





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