International IPPNW Congress 5 years living with Fukushima 30 years living with Chernobyl Berlin, February 27<sup>th</sup>, 2016 9.00 am – 10.30 am

## Thyroid Cancer under Age 19 in Fukushima

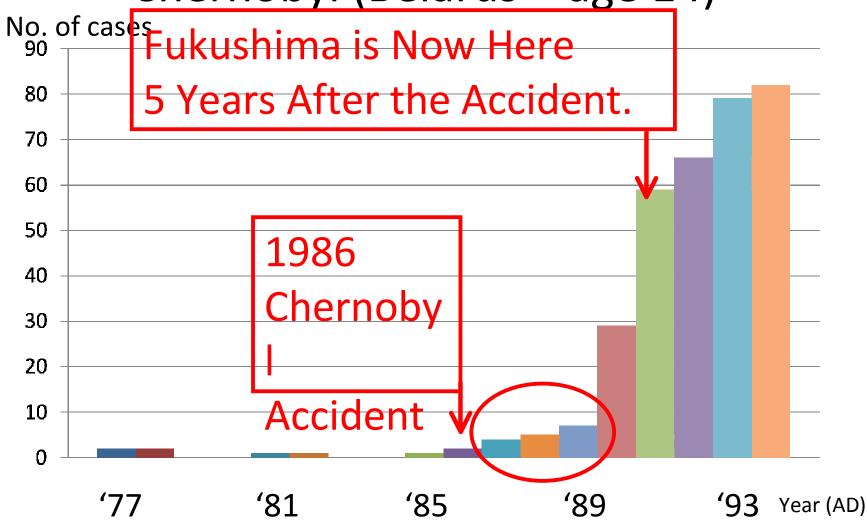
-As of 57 months after the accident-

(Analysis of data up to December 31, 2015, released on February 15, 2016 by Fukushima Prefecture)

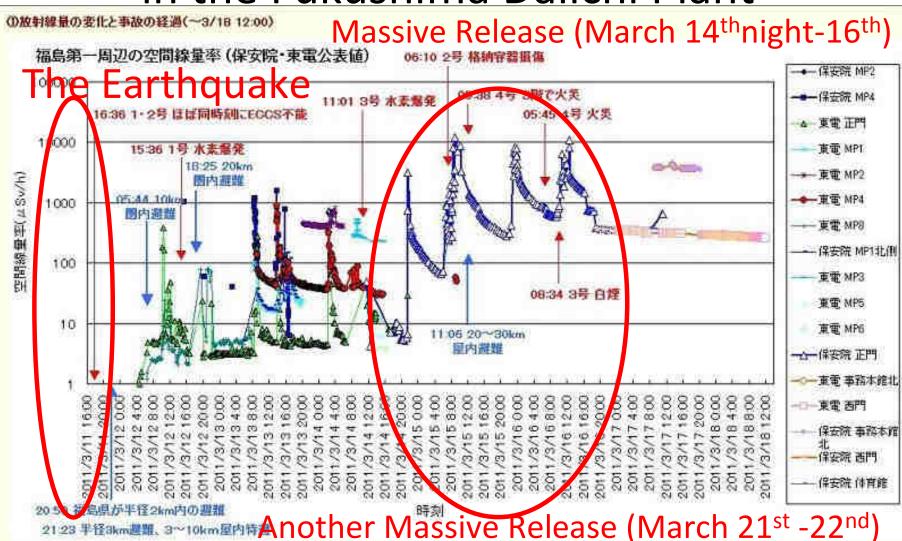
Toshihide Tsuda, Akiko Tokinobu, Etsuji Suzuki (Okayama University)

Eiji Yamamoto (Okayama University of Science)

# Epidemic Curve of Thyroid Cancer in Chernobyl (Belarus ≤ age 14)

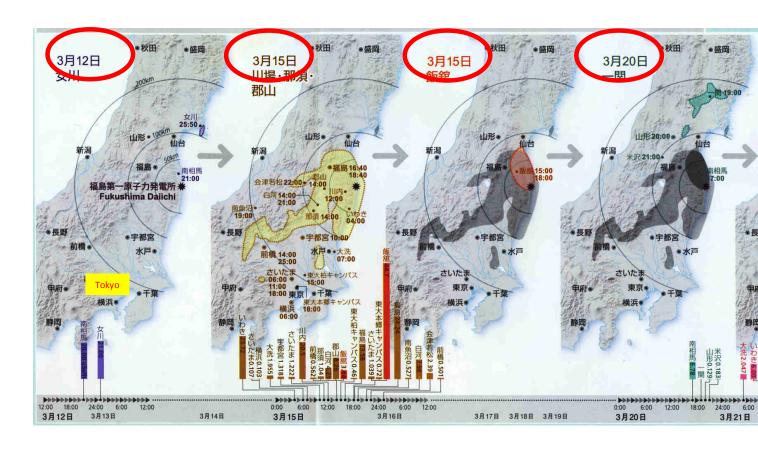


## Air dose rate after March 11, 2011 in the Fukushima Daiichi Plant



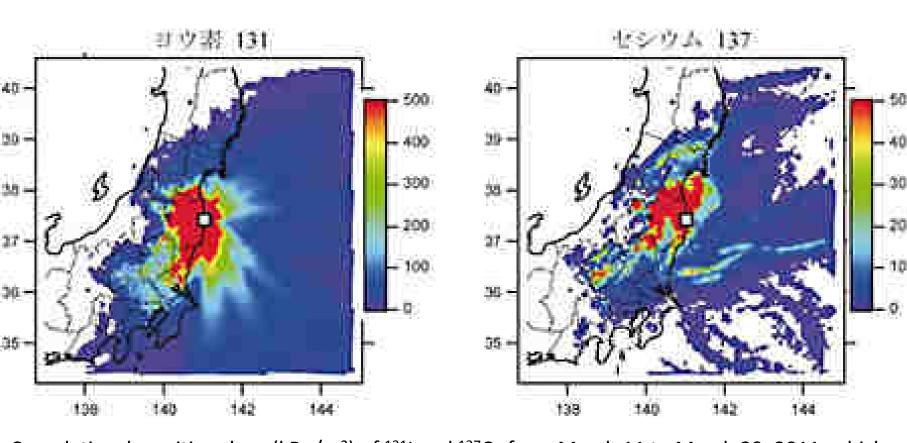
### Plume after March 11, 2011





By Prof. Y. Hayakawa Gunma University

### Different distributions of <sup>131</sup>I and <sup>137</sup>Cs



Cumulative deposition dose (kBq/m²) of <sup>131</sup>I and <sup>137</sup>Cs from March 11 to March 29, 2011, which estimated by Ohara T and Morino Y (National Institute for Environmental Studies): 「福島第一原子力発電所から放出された放射性物質の大気シミュレーション-東日本大震災復旧・復興への取組-」(October 31, 2011)

### Thyroid Cancer Screening ≤ age 18

- Primary examination: All residents ≤ age 18 in 2011 (born between April 2, 1992 and April 1, 2011)
  - Screened by thyroid ultrasound.
  - Secondary examination when nodules with diameter  $\geq$  5.1 mm or cysts with diameter  $\geq$  20.1mm are detected.
- Secondary examination: for positive primary examination
  - More detailed ultrasound, then cytology by fine needle aspiration.
- When cancer cells are detected by cytology
  - Followed with observation, then operated.
  - Cancer confirmed by histological examination of the excised tissues.

### Thyroid cancer screening schedule

- The First Round (The Japanese fiscal year is from April 1<sup>st</sup> to March 31<sup>st</sup> of the following year).
  - Year 1 (FY 2011, ending on March 31, 2012)
    - The nearest areas to the Fukushima Daiichi NPP
  - Year 2 (FY 2012, ending on March 31, 2013)
    - The moderately near areas including Fukushima City.
  - Year 3 (FY 2013, beginning on April 1, 2013)
    - Remaining areas ("Least Contaminated Area" by WHO 2012).
- The Second Round (from April 1,2014 to March 31, 2016)
  - Year 4 (FY 2014, ending on March 31, 2015)
    - The nearest areas and the moderately near areas
  - Year 5 (FY 2015, ending on March 31, 2016)
    - Remaining areas (corresponding to Year 3 in the first round)

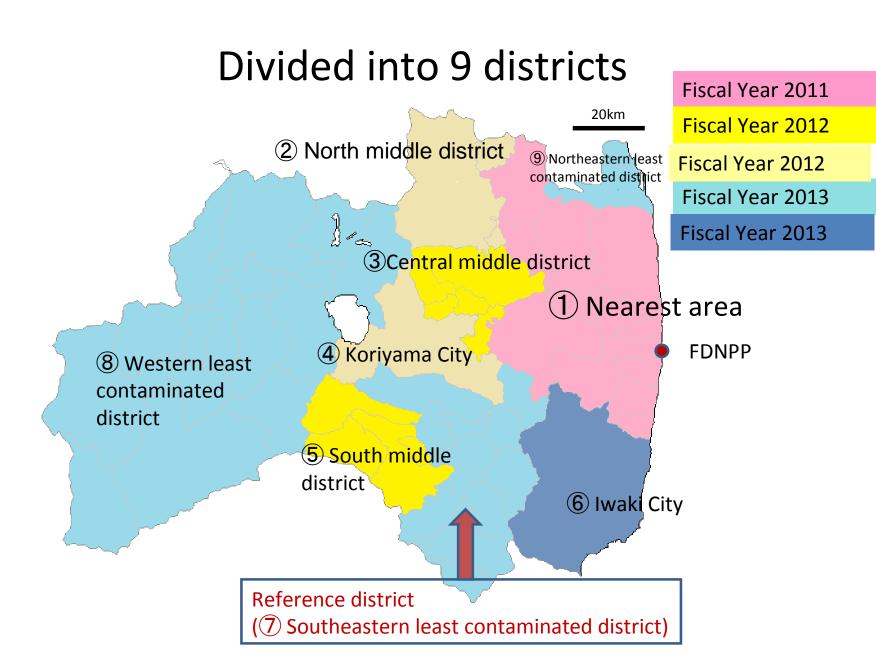
The order of screening according to the level of air dose rate



about three times higher than that of Gomel, Belarus.

#### Methods: Comparison Group

- Fukushima prefectural government releases the screening results about every three months.
  - The present data were released on February 15, 2016
     with the results as of December 31, 2015.
- Fukushima Prefecture was divided into 9 districts according to the screening schedule:
  - 1 Nearest area, 2 North middle district, 3 Central middle district, 4 Koriyama City, 5 South middle district, 6 Iwaki City, 7 Southeastern least contaminated district, 8 Western least contaminated district, 9 Northeastern least contaminated district



#### Methods: External Comparison

- Age- and sex-specific incidence estimates of thyroid cancer from the Center for Cancer Control and Information Services, National Cancer Center, Japan (1975-2008).
  - The Japanese mean annual incidence among those aged 0-19 years from 1975 to 2008 (i.e. 3 per 1,000,000) was used in the 1<sup>st</sup> round data. For the 2<sup>nd</sup> round data, we used 5 per 1,000,000 taking aging of the subjects into consideration.
- - In this case, "duration" is the duration from the date when thyroid cancer became detectable by screening and cytology to the date when it could have been diagnosed in usual clinical settings without screening.
- Poisson distribution was employed to estimate 95% confidence intervals.

#### Methods: Internal Comparison

- The southeastern least contaminated district was employed as "reference district."
- We estimated Prevalence Odds Ratio and its 95% Confidence Interval on the remaining 8 districts based on prevalence of the southeastern least contaminated district.
- We employed MLE Odds Ratio (Mid-P) of "StatCalc" in EpiInfo 7 (released by CDC).

#### Table 1

(1st round data: finalized as of June 30, 2015)

Areas	Population age ≤ 18	Participants in primary examination	Positives in primary examination	Participants in secondary examination	Thyroid Cancer Cases by FNAC(No. operated)	
Fiscal Year 2011	47,768	41,810 (87.5%)	221 (0.53%)	199 (90.0%)	15*(15*)	
Fiscal Year 2012	161,129	139,338 (86.5%)	988 (0.71%)	920 (93.1%)	56 (52)	
FY 2013 (Least Cont.)	158,788	119,328 (75.1%)	1,085 (0.91%)	989 (91.2%)	42 (32)	
Total	367,685	300,476 (81.7%)	2,294 (0.76%)	2,108 (91.9%)	113 (99) +3**(+2)**	

<sup>\*</sup>Including one benign case

n case \*\*Additional thyroid cancer cases released later

### Table 2 (4 districts in Middle Area)

Districts	Population age ≤ 18	Examinees in primary examination	Positives in primary examination	Examinees in secondary examination	Thyroid Cancer Cases by FNAC(No. operated)
North middle	57,211	50,618 (88.5%)	312 (0.62%)	298 (95.5%)	12 (?)
Central middle	21,052	18,194 (86.4%)	115 (0.63%)	111 (96.5%)	11 (?)
Koriyama City	64,380	54,063 (84.0%)	458 (0.85%)	415 (90.6%)	25 (?)
South middle	18,486	16,463 (89.1%)	103 (0.63%)	96 (93.2%)	8 (?)
Total	161,129	139,338 (86.5%)	988 (0.71%)	920 (93.1%)	56 (52)

Table 3:

4 Districts in the Least Contaminated Area

Participants Positives in Participants Thyroid

(0.85%)

(0.92%)

(0.81%)

(0.99%)

1,085

(0.91%)

455

242

334

(90.7%)

(92.7%)

(91.3%)

(88.9%)

(91.2%)

422

221

297

989

0(0)

24 (?)

7(?)

11 (?)

42 (32)

Districts	Population age ≤ 18	in primary	Positives in primary examination	in	Thyroid Cancer Cases by FNAC(No. operated)
Nicologic	0.246	6,359	54	49	0 (0)

(77.1%)

49,429

(79.3%)

29,820

(77.8%)

33,720

(67.5%)

119,328

(75.1%)

8,246

62,293

38,322

49,927

158,788

Northeastern

Southeastern

(Reference)

Western

Total

**Iwaki City** 

#### Table 4: External Comparison (1st Round)

	3/1,000,000*		Prevalence	
Areas and Districts	IRR**	(95% C.I.) **	per 10 <sup>6</sup>	Reciprocal
1 Nearest area (FY 2011)	29.90	(16.73-49.31)	359	2,787.3
2 North middle district	19.76	(10.21-34.51)	237	4,218.2
3 Central middle district	50.38	(25.15-90.15)	605	1,654.0
4 Koriyama City	38.54	(24.94-56.89)	462	2,162.5
5 South middle district	40.49	(17.48-79.79)	486	2,057.9
<b>6</b> Iwaki City	40.46	(25.92 -60.20)	486	2,059.5
⑦SE least contaminated district	19.56	(7.86-40.31)	235	4,260.0
8 Western least contaminated district	27.18	(13.57-48.64)	326	3,024.5
9NE least contaminated district	0	(0.00-48.34)	0	-
		//		. `

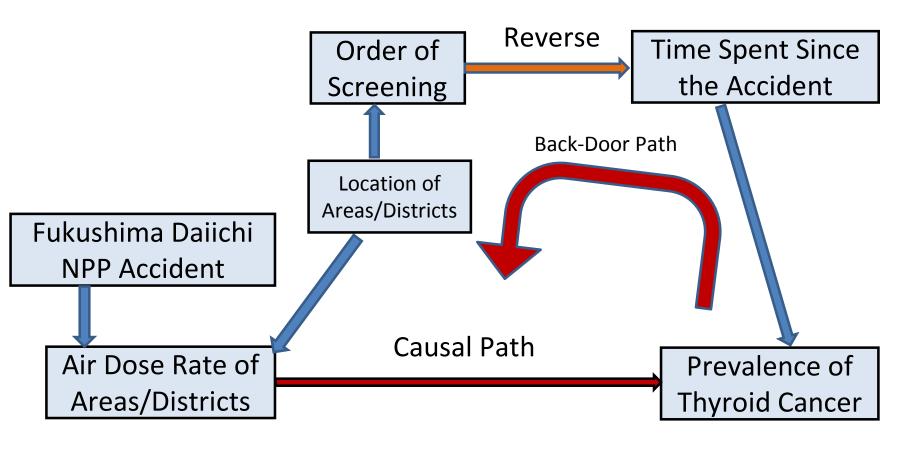
<sup>\*</sup>Comparison with Japanese mean \*\*Incident Rate Ratio (95% Confidence Interval)

Table 4-2: Internal Comparison and POR (1st round)

	Cancer cases	No. in primary exam.	POR*	(95% C.I.) *
①Nearest area (Fiscal Year 2011)	15**	41,810	1.53	(0.63-4.01)
2 North middle district	12	50,618	1.01	(0.40-2.73)
3 Central middle district	11	18,194	2.58	(0.99-7.06)
4 Koriyama City	25	54,063	1.97	(0.88-4.91)
5 South middle district	8	16,463	2.07	(0.73-6.00)
6 Iwaki City	24	49,429	2.07	(0.92-5.17)
7) SE least contaminated (Reference)	7	29,820	1	
8 Western least contaminated district	11	33,720	1.39	(0.54-3.81)
9NE least contaminated district	0	6,359	0	(0-2.50)

<sup>\*</sup> Prevalence Odds Ratio (95% Confidence Interval) \*\*Including one benign case

## Back-Door Path between Areas/Districts and Prevalence of Cancer



# Table 5 Latency Adjusted External (IRR) and Internal Comparison (POR)

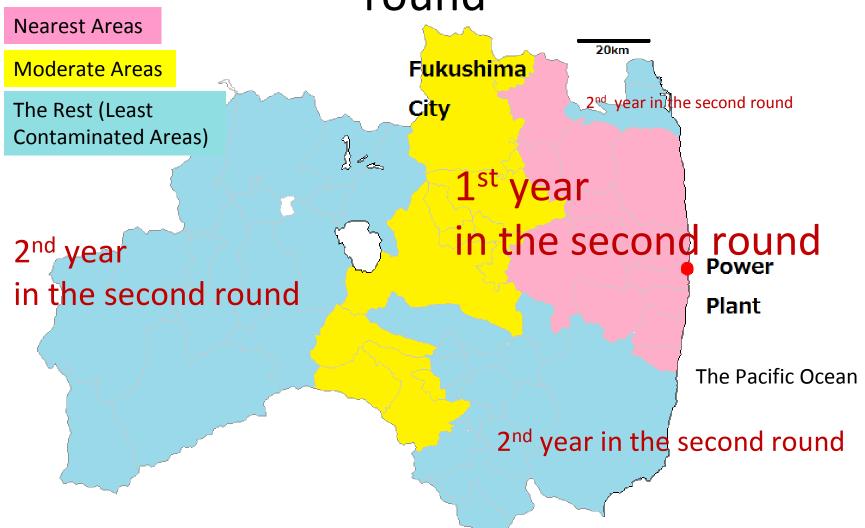
	Latency 1-3 years	Latency 2-4 years	
	IRR (95% CI) POR(9	5% CI) IRR (95% CI)	
1)2011 FY Combined (Nearest area)	120 (67 - 197) <mark>4.6 (2</mark> .	<b>1 – 11</b> ) 60 (33 - 99)	
2012 FY Combined (Middle area)	67 (51 - 87) 2.6 (1.	2 – 6.0) 45 (34 - 58)	
②North middle district	40 (20 - 69) <b>1.5 (0.</b>	<del>65 – 3.9)</del> 26 (14 - 46)	
③Central middle district	101 (50 - 180) 3.9 (1.	<mark>6 – 10)</mark> 67 (34 -120)	
Koriyama City district	77 (50 - 112) 3.0 (1.	<b>4 – 7.2)</b> 51 (33 - 76)	
⑤South middle district	81 (35 - 160) <b>3.1 (1</b> .	<b>2 – 8.4)</b> 54 (23 -106)	
2013 FY Combined (Least contaminated)	37 (26 - 50)	28 (20 - 38)	
6 Iwaki City district	50 (31 - 76) <b>2.1 (0.</b> 9	9 <mark>2 – 5.2)</mark> 38 (24 - 57)	
<b>⑦</b> Southeastern least contaminated	26 (11 - 54) 1 (refe	rence) 20 (8 – 41)	
8 Western least contaminated district	34 (16 - 62) 1.4 (0.5	5 <mark>4 - 3.8)</mark> 25 (12 - 47)	
Northeastern least contaminated	0 (0 - 66) 0 (0 -	<b>- 2.5)</b> 0 (0 - 50)	

Table 6: Internal Comparison and POR

	Cancer cases	No. in primary exam.	POR*	(95% C.I.) *
①Nearest area (Fiscal Year 2011)	15**	41,810	1.53	(0.63-4.01)
2 North middle district	12	50,618	1.01	(0.40-2.73)
3 Central middle district	11	18,194	2.58	(0.99-7.06)
4 Koriyama City	25	54,063	1.97	(0.88-4.91)
5 South middle district	8	16,463	2.07	(0.73-6.00)
6 Iwaki City	24	49,429	2.07	(0.92-5.17)
7)SE least contaminated (Reference)	7	29,820	1	
8 Western least contaminated district	11	33,720	1.39	(0.54-3.81)
9NE least contaminated district	0	6,359	0	(0-2.50)

<sup>\*</sup> Prevalence Odds Ratio (95% Confidence Interval) \*\*Including one benign case

The order of screening in the 2<sup>nd</sup> round



## Table 7: Current results of the 2<sup>nd</sup> round screening (as of December 31, 2015)

Areas	Population age* ≤ 18 (including fetuses)	Participants in primary exam.	Positives in primary exam.	Participants in secondary exam.	No. of FNAC	Thyroid Cancer Cases by FNAC(No. operated)
2014	216,874	155,536 (71.7%)	1,260 (0.8%)	990 (78.6%)	139	45 (?)
2015	164,387	81,059 (49.3%)	559 (0.7%)	182 (32.6%)	18	6 (?)
Total	381,261	236,595 (62.1%)	1,819 (0.8%)	1,172 (64.4%)	157	51 (16)

<sup>\*</sup> Age at the time of the March 2011 accident

Table 8. Results of Fiscal Vear 2011 in 2nd Round

Table 8: Results of Fiscal Year 2014 in 2114 Round							
(Data as of December 31, 2015)							
Districts/Area	Population age ≤ 18	Examinees in primary examination	Positives in primary examination	Examinees in secondary examination	Thyroid Cancer Cases by FNAC(No. operated)		
Nearest area	49,457	33,721 (66.6%)	332 (0.98%)	279 (84.0%)	16 (?)		
North middle	59,496	45,190 (75.6%)			9 ( 7 )		
Central middle	21,809	16,136 (73.9%)			4 (?)		
Koriyama City	66,759	45,965 (68.9%)			15 (7)		
South middle	19,353	14,524 (75.0%)			1 ( ? )		

155,536 1,260

(71.7%) (0.81%)

216,874

Total

#### Table 9: Results of FY 2015 in 2<sup>nd</sup> Round

4 Districts in the Least Contaminated Area (Data as of December 31, 2015)

(Data as of December 31, 2013)							
Districts	Population age ≤ 18	Participants in primary examination	Positives in primary examination	Participants in secondary examination	Thyroid Cancer Cases by FNAC(No. operated)		
Northeastern	8,563	5,463 (63.8%)	41 (0.75%)	28 (68.3%)	1 (?)		
Iwaki City	64,294	32,992 (51.3%)	277 (0.84%)	41 (14.8%)	2 (?)		
Southeastern (Reference)	39,772	26,665 (67.0%)	183 (0.69%)	98 (53.6%)	2 (?)		
Western	51,758	15,939 (30.8%)	58 (0.36%)	15 (25.9%)	1 (?)		
Total	164,387	46,865 (28.5%)	559 (1.19%)	182 (32.6%)	6 (?)		

### Table 10: External Comparison in the 2<sup>nd</sup> Round (Latency 2 years)

	5/1,000,000*		Prevalence	
Areas and Districts	IRR**	(95% C.I.) **	per 10 <sup>6</sup>	Reciprocal
1 Nearest area (Latency 2.5 years)	37.96	21.70-61.64)	474	2,107.6
2 North middle district	19.92	(9.11-37.81)	199	5,021.1
3 Central middle district	24.79	(6.75-64.46)	248	4,034.0
4 Koriyama City	32.63	(18.26-53.82)	326	3,064.3
5 South middle district	6.89	(0.17-38.36)	68.9	14,524
6 Iwaki City	6.06	(0.73-21.90)	60.6	16,496
7SE least contaminated district	7.50	(0.91-27.09)	75.0	13,332
®Western least contaminated district	6.27	(0.16-34.96)	62.7	15,939
9NE least contaminated district	18.30	(0.46-101.99)	183.1	5,463

#### **Tentative Conclusion 1**

- In Chernobyl, the outbreak of thyroid cancer actually began one year after the accident, rather than the fourth or fifth post-accident year when the marked increase was observed.
- As of 57 months after the accident, the first round screening conducted from October 2011 to March 2014 revealed a 20- to 50-fold excess incidence in thyroid cancer in ages 18 years or younger.
  - A higher rate of thyroid cancer with a dose-response tendency was seen with proximity to FDNPP, especially in the south area.

#### **Tentative Conclusion 2**

- Post-operative findings indicate that 92% of the operated cases had lymph node metastasis, distant metastasis and/or extrathyroidal extension.
- In the ongoing 2<sup>nd</sup> round screening, 20- to 38-fold excesses are already seen even though many results are still pending. Screening effect discussed in the 1<sup>st</sup> round has no relevance in the 2<sup>nd</sup> round. About 80% of thyroid cancer cases detected in the 2<sup>nd</sup> round had no lesions with pre-malignant potential detected in the 1<sup>st</sup> round: it appears that these cancers grew over 5.0 mm in diameter in only 2 years.
- We must prepare to take measures to deal with not only thyroid cancer but also other cancers and non-cancer diseases.
- Further investigation is needed especially in children in neighboring prefectures as well as in residents older than 18.

#### What is next?

- Fukushima thyroid cancer screening data is updated by Fukushima Prefecture every three months.
- Acknowledgement: We are grateful to Colin L. Soskolne, PhD, Martin Tondel, MD, PhD, Erik R. Svendsen, PhD, Gaston Meskens, MSc, and Wael Al-Delaimy, MD, PhD, for their thoughtful suggestions and constructive discussions on cancer-related issues relating to the 2011 nuclear accident in Fukushima, Japan. The authors also thank Tetsuji Imanaka, MSc, Keiji Hayashi, MD, and Okujou Iwami MD, PhD, for providing important references. We are also grateful to Professor Toshiro Ogura (Health Service Center of Okayama University) for providing their data, and Dr. Yuri Hiranuma for her thoughtful suggestions.
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