

Incidence trends of invasive and *in situ* breast cancer among females in Poland during 1999–2014

Trendy zachorowalności na inwazyjnego i przedinwazyjnego raka piersi u kobiet w Polsce w latach 1999–2014

Paweł Macek^{1,2}, Dana Hashim³, Marta Mańczuk⁴, Ewa Błaszczkiewicz^{1,5}, Barbara Sosnowska-Pasiarska⁶, Małgorzata Biskup^{7,8}, Halina Król^{7,9}, Jolanta Smok-Kalwat¹⁰, Stanisław Gózdź^{7,10}

¹Department of Epidemiology and Cancer Control, Holycross Cancer Centre, Kielce, Poland
Head of the Department: Paweł Macek PhD

²Faculty of Medical Sciences, School of Economics, Law, and Medical Sciences, Kielce, Poland
Head of the Faculty: Grzegorz Gatuszka PhD

³Department of Preventive Medicine, Institute of Translational Epidemiology, Icahn School of Medicine at Mount Sinai, One Gustave L. Levy Place, New York, USA
Head of the Department: Emanuela Taioli MD, PhD

⁴Cancer Primary Prevention Unit, Department of Cancer Epidemiology, The Maria Skłodowska-Curie Memorial Cancer Centre and Institute of Oncology, Warsaw, Poland
Head of the Unit: Marta Mańczuk PhD

⁵Department of Education, Jan Kochanowski University, Kielce, Poland
Head of the Department: Agnieszka Świerczek MA

⁶Department of Oncocardiology, Holycross Cancer Centre, Kielce, Poland
Head of the Department: Barbara Sosnowska-Pasiarska MD, PhD

⁷Faculty of Medicine and Health Sciences, Jan Kochanowski University, Kielce, Poland
Head of the Faculty: Prof. Stanisław Głuszek MD, PhD

⁸Department of Rehabilitation, Holycross Cancer Centre, Kielce, Poland
Head of the Department: Anna Opuchlik PhD

⁹Department of Didactics and Research, Holycross Cancer Centre, Kielce, Poland
Head of the Department: Halina Król PhD

¹⁰Clinic of Clinical Oncology, Holycross Cancer Centre, Kielce, Poland
Head of the Clinic: Stanisław Gózdź MD, PhD, Prof. UJK

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Słowa kluczowe: rak piersi, zachorowalność, epidemiologia, zdrowie publiczne.

Abstract

Introduction: Breast cancer is the most common malignant tumour among females worldwide, including low and middle-income countries. It affects 25% of the female population and yearly is diagnosed in 1.5 million women.

Aim of the research: To determine the trends and distributions for invasive and *in situ* breast cancer incidence rates between 1999 and 2014 in Poland overall and in the voivodeships with the greatest proportion of females aged ≥ 65 years old.

Material and methods: Data for 226,146 invasive (C50) and 7865 *in situ* (D05) breast cancer incidence (1999–2014) for all females was obtained from published data of the Polish National Cancer Registry (NCR). Age-standardised rates (ASRs) per 10^5 person-years for all ages of combined females with 95% confidence intervals (CI) were calculated. Joinpoint regression was performed to identify log-linear trends in both invasive and carcinoma *in situ* incidence rates in the analysed voivodeships and in Poland overall.

Results: The ASRs of both invasive and *in situ* breast cancers showed increasing trends over the 1999–2014 period, with different patterns between analysed voivodeships. On average, average annual percent changes (AAPCs) varied from 1.6% to 7.9% annually, while in the Świętokrzyskie Voivodeship AAPCs remained below 0.9% annually.

Conclusions: *In situ* and invasive breast cancer ASRs in Poland increased between 1999 and 2014. The Annual Percentage Change have slowed since 2004 in the analysed voivodeships, but not in Poland overall. It remains uncertain if breast cancer incidence rates will continue to increase or stabilise over time.

Streszczenie

Wprowadzenie: Rak piersi jest najczęściej występującym nowotworem złośliwym u kobiet na świecie. Dotyczy 25% populacji kobiet. Każdego roku diagnozuje się 1,5 miliona nowych zachorowań.

Cel pracy: Określenie tendencji czasowych zachorowalności na inwazyjnego i przedinwazyjnego raka piersi w latach 1999–2014 w Polsce oraz w województwach o najwyższym odsetku kobiet w wieku ≥ 65 lat.

Materiał i metody: Dane o 226 146 zachorowaniach na inwazyjnego (C50) i 7856 zachorowaniach na przedinwazyjnego (D05) raka piersi pochodziły z Krajowego Rejestru Nowotworów (KRN). Obliczono standaryzowane wg wieku współczynniki zachorowalności (ASR) wyrażone na 10^5 osobolat z 95-procentowym przedziałem ufności (CI). Zastosowano regresję Joinpoint w celu określenia tendencji log-liniowych trendów zachorowalności na inwazyjnego i przedinwazyjnego raka piersi w analizowanych województwach i w Polsce.

Wyniki: W latach 1999–2014 ASR dla inwazyjnego i przedinwazyjnego raka piersi wykazywały tendencję wzrostową ze zmiennymi wzorcami zachorowalności w analizowanych województwach. Średnia roczna zmiana procentowa (AAPC) wahała się od 1,6% do 7,9%, natomiast w województwie świętokrzyskim pozostawała na poziomie poniżej 0,9%.

Wnioski: W Polsce w latach 1999–2014 trendy czasowe zachorowalności na inwazyjnego i przedinwazyjnego raka piersi wykazywały tendencję wzrostową. W analizowanych województwach roczne zmiany procentowe zwolniły od 2004 r. Tendencji tej nie obserwowano w Polsce ogółem. Obserwowane zmiany estymatorów parametrów populacyjnych nie pozwalają na określenie wzorców zachorowalności na raka piersi w najbliższych latach.

Introduction

Breast cancer is the most common malignant tumour among females worldwide, including low and middle-income countries [1]. It affects 25% of the female population and was diagnosed in 1,676,633 women, age standardised rate (ASR): $43.3/10^5$ worldwide in 2012 [2]. Breast cancer incidence differs in various regions of the world, ranging from $26.8/10^5$ (10,922 new cases) in Central Africa to $96.0/10^5$ (161,529 new cases) in Western Europe. The breast cancer incidence rate in Poland was $51.9/10^5$ (17,259 diagnosed cases) for 2012, placing it among the medium-risk countries. Although Polish age-standardised incidence rates were about 1.7-times lower than in countries in western Europe, incidence rates were higher than in Central and Eastern European neighbouring countries [3].

After the Population Breast Cancer Early Detection Program was implemented in 2006, breast cancer screening programs in Poland were established nation-wide [4]. Most likely, as a result of increased screening, there has since been a reported increase in the number of ductal carcinoma *in situ* (DCIS) [5] and invasive early-stage breast cancer cases. Given that the comparison of 5-year relative survival rate values is often used as an indicator of early detection and treatment efficacy in breast cancer rate reduction [5], EURO CARE 5 data estimated the 5-year survival relative rate for women at 81.8% for Europe, but 71.6% for Poland (2000–2007) [6].

Poland is divided into 16 principal administrative regions (called voivodeships) and has an estimated population of 38.5 million (2013) [7]. In recent decades, there have been significant changes in the age structure of the Polish population; the population aged 65 and above increased to comprise approximately 15% (5.7 million) of the total population for the year 2013. Of those individuals age 65 and above, nearly 3.5 million were women (61%) [7]. This new age distribution implies a rising number of cancer-re-

lated deaths as well as substantially increased cancer care costs and complexity of care in the coming years.

Aim of the research

Given a lower 5-year survival rate compared to other European countries, the aim of this study was to determine the trends and distributions for invasive and *in situ* breast cancer incidence rates between 1999 and 2014 in Poland overall and in voivodeships with the greatest proportion of females aged ≥ 65 years old.

Material and methods

Study area and study design

Breast cancer incidence rates (1999–2014) were calculated among women living in voivodeships with the greatest proportion of females aged ≥ 65 years old. At least 19% of the women in the aforementioned voivodeships were over 65 years old. The percentage of this subpopulation for Poland overall is 17.6%. A total of 7 selected voivodeships was included (Figure 1) [8].

Data sources

Data for invasive (C50) and *in situ* (D05) breast cancer incidence (1999–2014) for all females was obtained from published data of the Polish National Cancer Registry (NCR) [9]. The analysis included information on invasive (C50) and *in situ* (D05) breast cancer cases coded according to the International Statistical Classification of Diseases and Related Health Problems ICD-10. The data regarding the population number within voivodeships and Poland overall were determined by the current place of residence, in all ages combined, and in 18 5-year age groups, starting from the age of 0 to 85 years and older, was also obtained from the NCR in Poland.

Ethics statement

This study is based on secondary data from the NCR. All of the presented information has been of-

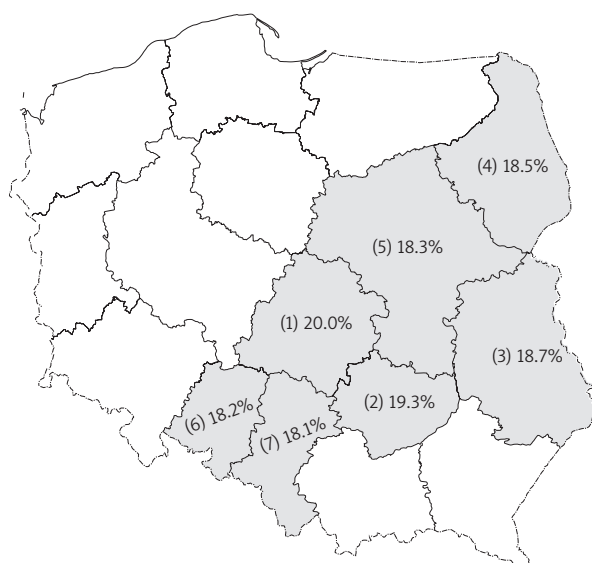


Figure 1. Voivodeships selected for incidence trend analysis were those in which the percentage of the female population ≥ 65 years old is over 18% (1 – Łódź, 2 – Świętokrzyskie, 3 – Lublin, 4 – Podlasie, 5 – Mazovia, 6 – Opole, 7 – Silesia; the percentage of females ≥ 65 years old for the whole of Poland is 17.6%) [8]

officially published on the NCR website and patients' consent was not required for accessing data because all data was anonymised.

Statistical analysis

Age-standardised rates (ASRs) per 100,000 person-years for all ages combining females with 95% confidence intervals (CI) for both invasive cancer and breast cancer *in situ* were calculated [10]. Standardisation was performed according to the age distribution of the world standardised population using the direct method [11]. Joinpoint regression was performed to identify statistically significant log-linear trends in both invasive and carcinoma *in situ* incidence rates in each voivodeship and in Poland overall. This method uses Monte Carlo permutation tests, with Bonferroni adjustment for multiple comparisons of significant joinpoints [12]. The Annual Percentage Change (APC) within each joinpoint segment was estimated with corresponding 95% confidence intervals (CI). The slope of the trend line corresponds to the value and direction of the APC incidence rates. For incidence trends in which more than one slope was identified, the average annual percent change (AAPC) from 1999 to 2014 (when available) was calculated based on the geometric mean of the APC trends [13] under the assumption of a constant rate of change. When this assumption does not hold over the entire time interval, the trend may be characterized using the annual percent changes from segmented analysis (sAPCs).

Although some breakpoints contained statistically significant nonlinear changes in trend slopes with

one to three joinpoints, most trends were linear with the smallest number of breakpoints equal to zero. Trend indication and APC assessment enabled the determination of the speed of breast cancer incidence changes among women (Tables 1, 2). In order to avoid an autocorrelation error, we decided to “fit model uncorrelated errors”. The program assumes that the random errors in the regression model were correlated and the regression coefficients estimated by ordinary least squares. From the trend analysis of *in situ* breast cancer, we excluded years of diagnosis with no cases of incidence.

Statistical analyses and graphic representations were conducted using Microsoft Office 2013, SAS Enterprise Guide version 7.1, and Joinpoint Regression Program version 4.0 (available from NCI, Bethesda, MD) [12].

Results

During 1999–2014, 234,011 cases of breast cancer were diagnosed in Polish women (226,146 – invasive breast cancer and 7865 – *in situ* breast cancer). Seven out of a total of 16 voivodeships were analysed with the number of cases totalling 108,777 (105,064 – invasive breast cancer and 3713 – *in situ* breast cancer). These constituted 46.4% of the total number of new breast cancer cases during 1999–2014 in Poland (46.4% of the total invasive breast cancers and 40.3% of *in situ* breast cancers in Poland).

Incident cases of both invasive and *in situ* breast cancer increased in the voivodeships with a high proportion of women aged ≥ 65 years and in Poland overall. In 1999, the prevalence of *in situ* breast cancer varied from 0.0% (in Łódź, Lublin, Opole, Podlasie) to 2.5% (in Mazovia). Between 1999–2005 the ratio of carcinoma *in situ* versus invasive cancers increased to 6.8% (in 2005 in Świętokrzyskie). From 2006 onwards the ratio of *in situ* vs. invasive breast cancer increased to 8.6% (in 2014 in Świętokrzyskie). This suggests an effect of early detection by the screening program implemented in Poland from 2007. Within the total time period (1999–2014) the ratio of *in situ* vs. invasive breast cancer ranged from 1.2% (in Łódź) to 5.8% (in Świętokrzyskie), with an average of 3.3% (not shown).

The ASR of invasive breast cancer among Polish women ranged from 42.1 in 1999 to 55.8 in 2014. Among women in analysed voivodeships and by year, the lowest invasive breast cancer ASRs were 18.1 (in 1999, Łódź Voivodeship) and highest 60.9 (in 2014, Łódź Voivodeship) (Table 3).

The ASRs of *in situ* breast cancer among women in Poland ranged from 0.3 (in 1999) to 3.4 (in 2014). Among women in analysed voivodeships, the lowest ASRs were 0.0 per 100,000 (in several years in various voivodeships) and the highest was 4.4/100,000 (in 2014, Świętokrzyskie Voivodeship) (Table 4).

Table 1. Joinpoint regression analysis of incidence of invasive (C50) breast cancer among women in selected voivodeships and in Poland overall, 1999–2014

Voivodeship	Trend 1		Trend 2		Trend 3		Full range
	Years	APC (95% CI)	Years	APC (95% CI)	Years	APC (95% CI)	AAPC (95% CI)
Świętokrzyskie	1999–2001	4.4 (–9.0, 19.8)	2001–2004	–6.0 (–16.7, 6.1)	2004–2014	2.4 [^] (1.8, 2.9)	0.9 (–1.7, 3.5)
Mazovia	1999–2014	1.9 [^] (1.5, 2.3)	–	–	–	–	1.9 [^] (1.5, 2.3)
Silesia	1999–2014	1.6 [^] (1.0, 2.1)	–	–	–	–	1.6 [^] (1.0, 2.1)
Łódź	1999–2001	31.1 [^] (4.9, 63.8)	2001–2006	9.4 [^] (4.1, 14.9)	2006–2014	1.8 [^] (0.2, 3.4)	7.9 [^] (4.7, 11.1)
Lublin	1999–2001	24.4 (–9.3, 70.5)	2001–2014	1.8 [^] (0.7, 2.8)	–	–	4.5 [^] (0.6, 8.6)
Opole	1999–2014	2.1 [^] (1.4, 2.7)	–	–	–	–	2.1 [^] (1.4, 2.7)
Podlasie	1999–2014	1.6 [^] (0.9, 2.4)	–	–	–	–	1.6 [^] (0.9, 2.4)
Poland	1999–2014	2.0 [^] (1.7, 2.4)	–	–	–	–	2.0 [^] (1.7, 2.4)

APC – Annual Percentage Change, AAPC – Average Annual Percentage Change, CI – confidence interval, [^](APC/AAPC) is significantly different from zero at $\alpha = 0.05$; 95% CI (APC/AAPC) is corresponding with 95% confidence intervals.

Table 2. Joinpoint regression analysis of incidence of *in situ* (D05) breast cancer among women in selected voivodeships and in Poland overall, 1999–2014

Voivodeship	Trend 1		Trend 2		Trend 3		Full range
	Years	APC (95% CI)	Years	APC (95% CI)	Years	APC (95% CI)	AAPC (95% CI)
Świętokrzyskie	2001–2004	114.8 [^] (34.7, 242.6)	2004–2014	3.9 (–1.5, 9.5)	–	–	22.8 [^] (11.2, 35.7)
Mazovia	1999–2014	5.3 [^] (3.8, 7.0)	–	–	–	–	5.3 [^] (3.8, 7.0)
Silesia	1999–2002	–7.2 (–21.8, 10.1)	2002–2014	17.2 [^] (15.5, 18.9)	–	–	11.9 [^] (8.3, 15.5)
Łódź	2004–2014	8.2 [^] (4.9, 11.6)	–	–	–	–	8.2 [^] (4.9, 11.6)
Lublin	2002–2004	186.2 (–11.6, 826.4)	2004–2014	2.8 (–2.3, 8.3)	–	–	22.0 [^] (2.9, 44.6)
Opole	2003–2008	17.7 [^] (8.5, 27.7)	2008–2011	–17.9 (–48.8, 31.5)	2011–2014	17.0 (–4.6, 43.5)	6.5 (–3.9, 18.0)
Podlasie	2000–2008	17.6 [^] (7.2, 29.0)	2008–2011	–33.0 (–83.5, 171.2)	2011–2014	56.5 (–10.8, 174.5)	10.8 (–15.5, 45.4)
Poland	1999–2001	53.6 [^] (19.6, 97.4)	2001–2010	15.3 [^] (13.7, 16.9)	2010–2014	5.8 [^] (0.9, 11.0)	17.1 [^] (13.5, 20.8)

APC – Annual Percentage Change, AAPC – Average Annual Percentage Change, CI – confidence interval; [^](APC/AAPC) is significantly different from zero at $\alpha = 0.05$; 95% CI (APC/AAPC) is corresponding with 95% confidence intervals.

The age-standardised incidence rates of both invasive and *in situ* breast cancers showed increasing trends during the 1999–2014 period, with different patterns between analysed voivodeships (Tables 1, 2).

Time trends in invasive breast cancer incidence between 1999 and 2014 have varied slightly across the analysed data sources. The results of analyses indicated that most incidence rates increased within voivodeships and at the national level. On average, AAPCs varied from 1.6% to 7.9% annually, while in

Świętokrzyskie voivodeship AAPCs were remained below 0.9% annually (Figure 2).

For invasive breast cancer, ASRs for the 1999–2014 period, an increase (with no joinpoints) was observed in Podlasie, Mazovia, Opole, and Silesia voivodeships and at the national level in Poland (by 1.6% to 2.1% per year). In one voivodeship, there was one joinpoint – in Lublin (in 2001). In Lublin voivodeship, the ASRs increased by 24.4% during 1999–2001 and by 1.8% per year between 2001 and 2014. In Świętokrzyskie

Table 3. Age-standardized incidence rates of invasive (C50) breast cancer among women in selected voivodeships and in Poland, 1999–2014

Voivodeship	Year of diagnosis																
	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	1999–2014
Świętokrzyskie	N	410	405	433	444	395	370	406	398	411	417	426	472	472	535	510	6971
	ASR	44.6	45.5	46.3	47.1	42.5	38.3	41.2	42.0	43.3	42.5	42.6	46.7	46.8	51.7	48.9	44.8
	95% CI	(40.3, 48.9)	(41.1, 50.0)	(42.0, 50.7)	(42.7, 51.4)	(38.4, 46.7)	(34.4, 42.2)	(37.2, 45.2)	(46.1, 46.1)	(37.9, 47.5)	(39.1, 47.5)	(38.4, 46.6)	(38.6, 46.7)	(42.4, 50.9)	(42.6, 51.0)	(47.4, 56.1)	(44.7, 53.2)
Mazovia	N	1594	1608	1605	1731	1565	1731	1767	1823	2064	2007	2271	2102	2162	2447	2315	31147
	ASR	44.3	44.6	43.8	46.7	41.8	45.7	45.7	47.0	52.1	50.1	55.7	50.9	51.9	57.7	54.1	49.5
	95% CI	(42.2, 46.5)	(42.4, 46.8)	(41.7, 46.0)	(44.5, 48.9)	(39.7, 43.8)	(43.5, 47.8)	(43.6, 47.8)	(44.8, 49.1)	(44.8, 49.1)	(49.9, 54.4)	(47.9, 52.3)	(48.7, 53.0)	(49.7, 54.1)	(55.4, 60.0)	(51.9, 56.3)	(52.6, 57.0)
Silesia	N	1532	1525	1471	1459	1478	1473	1754	1667	1682	1634	1869	1834	2118	2022	2013	27565
	ASR	45.7	45.0	43.3	42.9	42.7	42.2	49.3	46.4	46.6	44.9	50.6	49.1	56.4	53.0	52.2	47.8
	95% CI	(43.4, 48.0)	(42.8, 47.3)	(41.1, 45.5)	(40.7, 45.1)	(40.6, 44.9)	(40.0, 44.3)	(47.0, 51.6)	(44.1, 48.6)	(44.4, 48.8)	(42.7, 47.1)	(48.3, 52.9)	(46.9, 51.3)	(54.0, 58.9)	(50.7, 55.3)	(49.9, 54.4)	(50.2, 54.7)
Łódź	N	360	583	637	764	702	926	1098	1064	1080	1162	1248	1253	1188	1408	1303	16155
	ASR	18.1	29.8	32.6	39.3	36.2	46.3	53.5	51.6	52.3	55.5	58.7	59.7	57.5	62.4	59.1	48.8
	95% CI	(16.3, 20.0)	(27.3, 32.2)	(30.1, 35.1)	(36.6, 42.1)	(33.5, 38.9)	(43.4, 49.3)	(50.3, 56.6)	(48.5, 54.7)	(49.1, 55.4)	(52.3, 58.6)	(55.4, 61.9)	(56.4, 63.0)	(54.2, 60.7)	(59.2, 65.7)	(55.9, 62.3)	(48.0, 49.5)
Lublin	N	365	542	594	647	606	585	856	722	735	684	772	848	778	813	895	844
	ASR	24.6	36.4	38.4	43.2	39.5	38.0	54.1	44.8	45.7	42.3	47.6	51.3	47.1	48.9	51.7	48.9
	95% CI	(22.1, 27.1)	(33.3, 39.4)	(35.3, 41.5)	(39.9, 46.6)	(36.3, 42.6)	(35.0, 41.1)	(50.4, 57.7)	(41.5, 48.1)	(42.4, 49.0)	(39.1, 45.5)	(44.3, 51.0)	(47.8, 54.8)	(43.8, 50.4)	(45.5, 52.2)	(48.3, 55.1)	(45.6, 52.2)
Opole	N	271	304	318	318	279	330	328	339	394	390	410	376	449	435	449	5814
	ASR	38.2	41.8	43.9	42.8	37.0	42.7	42.7	45.1	49.9	48.9	51.0	45.6	54.1	52.4	52.5	46.3
	95% CI	(33.7, 42.8)	(37.1, 46.5)	(39.1, 48.7)	(38.1, 47.5)	(32.6, 41.3)	(38.1, 47.3)	(38.0, 47.3)	(40.3, 49.9)	(40.3, 49.9)	(44.9, 54.8)	(46.1, 56.0)	(41.0, 50.2)	(49.1, 59.2)	(47.5, 57.3)	(47.6, 57.3)	(45.1, 54.1)
Podlasie	N	345	295	339	370	347	302	368	354	368	413	408	410	435	434	469	6126
	ASR	43.0	37.5	42.6	46.2	42.3	35.7	43.6	41.8	43.7	48.9	48.0	46.0	47.8	49.1	50.4	45.0
	95% CI	(38.5, 47.6)	(33.2, 41.8)	(38.0, 47.1)	(41.5, 50.9)	(37.8, 46.8)	(31.7, 39.7)	(39.2, 48.1)	(37.4, 46.1)	(39.3, 48.2)	(44.2, 53.6)	(43.4, 52.7)	(41.6, 50.5)	(43.3, 52.3)	(44.5, 53.7)	(45.8, 54.9)	(46.3, 55.5)
Poland	N	10903	11853	12118	12132	11733	12049	13385	13322	14484	14576	15752	15784	16534	17000	17142	226146
	ASR	42.1	45.4	45.9	45.5	43.6	44.0	48.1	47.6	51.1	50.7	54.2	53.4	55.6	56.0	55.8	49.9
	95% CI	(41.3, 42.9)	(44.6, 46.3)	(45.1, 46.7)	(44.7, 46.4)	(42.8, 44.4)	(43.2, 44.8)	(47.3, 48.9)	(46.8, 48.4)	(50.3, 52.0)	(49.9, 51.5)	(53.4, 55.1)	(52.5, 54.2)	(54.8, 56.5)	(55.2, 56.9)	(55.0, 56.7)	(49.7, 50.1)

ASR – age-standardized rate, N – numbers of new breast cancer cases; 95% CI – 95% confidence interval, ASR per 100,000 (direct method, world standard population).

Table 4. Age-standardized incidence rates of *in situ* (D05) breast cancer among women in selected voivodeships and in Poland, 1999–2014

Voivodeship	Year of diagnosis																	
	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	1999–2014	
Święto-krzyskie	N	2	0	1	15	8	23	27	36	33	35	33	26	56	36	44	402	
	ASR	0.3	0.0	0.1	1.7	1.0	2.6	2.9	3.8	3.4	3.7	3.4	2.7	6.1	3.5	4.4	2.7	
	95% CI	(-0.1, 0.6)	nc	(-0.1, 0.4)	(0.9, 2.6)	(0.3, 1.7)	(1.5, 3.6)	(2.0, 4.4)	(1.8, 4.0)	(2.5, 5.0)	(2.2, 4.5)	(2.5, 4.9)	(2.3, 4.6)	(1.7, 3.8)	(4.5, 7.7)	(2.3, 4.6)	(3.1, 5.7)	(2.5, 3.0)
Mazovia	N	40	60	72	100	67	72	100	84	114	108	100	150	139	121	137	1574	
	ASR	1.2	1.7	2.0	2.8	1.8	1.9	2.7	2.4	2.9	2.7	2.8	3.7	3.4	3.2	3.4	2.6	
	95% CI	(0.8, 1.6)	(1.3, 2.1)	(1.5, 2.4)	(2.3, 3.4)	(1.4, 2.3)	(1.5, 2.4)	(2.2, 3.3)	(1.9, 2.9)	(2.4, 3.5)	(2.2, 3.2)	(2.3, 3.3)	(2.0, 3.0)	(3.1, 4.3)	(2.9, 4.0)	(2.6, 3.7)	(2.8, 3.9)	(2.5, 2.7)
Silesia	N	24	27	20	21	28	18	47	36	53	56	79	102	123	141	128	974	
	ASR	0.7	0.8	0.7	0.6	0.9	0.6	1.3	1.0	1.5	1.6	2.1	2.8	3.3	3.7	3.4	1.7	
	95% CI	(0.4, 1.0)	(0.5, 1.1)	(0.4, 1.0)	(0.4, 0.9)	(0.5, 1.2)	(0.3, 0.8)	(0.9, 1.7)	(0.7, 1.4)	(1.1, 1.9)	(1.2, 2.0)	(1.6, 2.6)	(2.2, 3.3)	(2.7, 3.9)	(3.1, 4.3)	(2.8, 4.0)	(1.6, 1.8)	
Łódź	N	0	0	0	1	0	13	13	13	11	17	29	12	27	18	31	197	
	ASR	0.0	0.0	0.0	0.0	0.0	0.6	0.7	0.6	0.6	0.8	1.5	0.6	1.4	1.0	1.4	0.6	
	95% CI	nc	nc	nc	(0.0, 0.1)	nc	(0.3, 1.0)	(0.3, 1.1)	(0.3, 1.0)	(0.2, 0.9)	(0.4, 1.2)	(0.9, 2.0)	(0.2, 0.9)	(0.8, 1.9)	(0.5, 1.4)	(0.9, 1.9)	(0.5, 0.7)	
Lublin	N	0	0	0	1	10	15	20	18	14	43	19	19	24	26	37	264	
	ASR	0.0	0.0	0.0	0.1	0.7	1.1	1.4	1.1	1.3	1.0	2.8	1.2	1.5	1.5	2.2	1.1	
	95% CI	nc	nc	nc	(-0.1, 0.2)	(0.3, 1.1)	(0.5, 1.6)	(0.8, 2.0)	(0.6, 1.7)	(0.7, 1.8)	(0.5, 1.6)	(1.9, 3.6)	(0.7, 1.7)	(0.9, 2.1)	(0.9, 2.0)	(1.5, 2.9)	(0.9, 1.2)	
Opole	N	0	0	0	0	11	12	15	17	24	25	15	19	16	23	21	216	
	ASR	0.0	0.0	0.0	0.0	1.6	1.5	2.0	2.4	3.1	3.4	2.4	2.1	2.0	2.8	2.6	1.8	
	95% CI	nc	nc	nc	nc	(0.7, 2.6)	(0.7, 2.4)	(1.0, 3.0)	(1.2, 3.5)	(1.8, 4.3)	(2.1, 4.8)	(0.9, 2.9)	(1.2, 3.1)	(1.0, 2.9)	(1.7, 4.0)	(1.5, 3.8)	(1.5, 2.0)	
Podlasie	N	0	2	2	5	2	8	5	10	4	7	5	3	5	8	11	86	
	ASR	0.0	0.3	0.3	0.7	0.3	1.0	0.7	1.5	0.5	1.2	0.9	0.3	0.6	0.8	1.5	0.7	
	95% CI	nc	(-0.1, 0.6)	(-0.1, 0.7)	(0.1, 1.3)	(-0.1, 0.7)	(0.3, 1.7)	(0.1, 1.4)	(0.6, 2.4)	(0.0, 1.1)	(0.4, 2.0)	(0.2, 1.6)	(0.0, 1.2)	(0.1, 0.7)	(0.1, 1.2)	(0.3, 1.4)	(0.6, 0.9)	
Poland	N	80	126	172	220	252	259	375	372	495	514	700	723	799	886	999	7865	
	ASR	0.3	0.5	0.7	0.9	1.0	1.0	1.4	1.4	1.8	1.9	2.5	2.5	2.8	3.1	3.0	3.4	1.8
	95% CI	(0.3, 0.4)	(0.4, 0.6)	(0.6, 0.8)	(0.8, 1.0)	(0.9, 1.1)	(0.8, 1.1)	(1.3, 1.6)	(1.2, 1.5)	(1.7, 2.0)	(1.7, 2.0)	(2.3, 2.7)	(2.3, 2.7)	(2.6, 3.0)	(2.9, 3.3)	(2.8, 3.2)	(3.2, 3.6)	(1.8, 1.8)

ASR – age-standardized rate, N – numbers of new breast cancer cases, 95% CI – 95% confidence interval, nc – not calculated, ASR per 100,000 (direct method, world standard population).

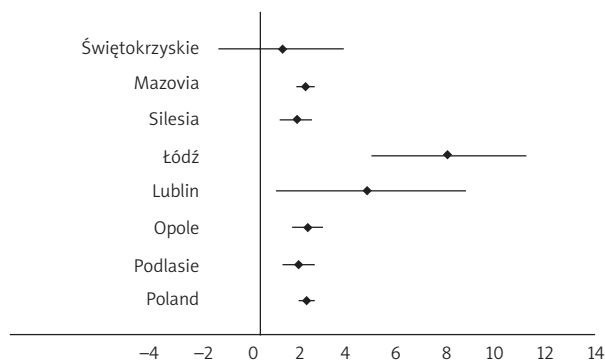


Figure 2. Average Annual Percentage Change (AAPC) of incidence of invasive breast cancer (C50) among women in selected voivodeships and in Poland, 1999–2014

voivodeship, there were two joinpoints – in 2001 and 2004. The ASRs increased by 4.4% between 1999 and 2001, decreased by 6.0% during 2001–2004, and then increased by 2.4% per year between 2004 and 2014. Two joinpoints (three trends) were also found in Łódź Voivodeship – in 2001 and 2006. The ASRs of invasive breast cancer increased by 31.1% per year during 1999–2001. From 2001 to 2006 and 2006 to 2014 the ASRs also increased (by 9.4% and 1.8% per year, respectively).

Time trends in *in situ* breast cancer incidence among women during the study period (1999–2014) increased on average (AAPCs varied from 5.3% in Mazovia to 22.8% in Świętokrzyskie voivodeship annually) and these changes were more dynamic than in the case of invasive breast cancer incidence trends (Figure 3).

The ASRs over the analysed period of time for *in situ* breast cancer and an increase (with no joinpoints) was observed only in Mazovia voivodeship (APC: 5.3%; period of time 16 years) and Łódź voivodeship (APC: 8.2%; period of time 11 years). Joinpoint analyses identified one joinpoint for the incidence series, separating two trends, in the rest of the voivodeships. In Świętokrzyskie (period of time 14 years) the ASRs of incidence increased by 114.8% per year between 2001 and 2004 (95% CI: 34.7–242.6), followed by an increasing trend from 2004 to 2014 by 3.9% per year.

In Silesia (period of time 16 years) the ASRs decreased from 1999 to 2002, and then they rapidly increased with a growing trend from 2002 to 2014 (APC: 17.2%).

In Lublin voivodeship (period of time 13 years) the values of age-standardised incidence rates increased by 186.2% per year from 2002 to 2004, and from 2004 to 2014 the trend of increase was 2.8% per year.

In Podlasie voivodeship (period of time 15 years) the ASRs increased by 17.6% from 2000 to 2008, decreased by 33.0% from 2008 to 2011, then again they rapidly increased by 56.5% per year from 2011 to

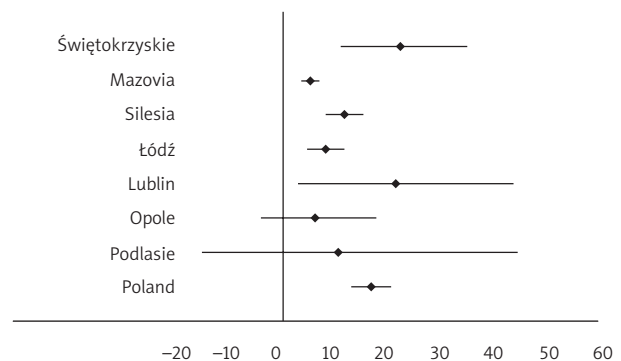


Figure 3. Average Annual Percentage Change (AAPC) of incidence of in situ breast cancer (D05) among women in selected voivodeships and in Poland in the analysed periods

2014. Two joinpoints were found at the national level (in 2001 and in 2010).

Opole voivodeship (period of time 12 years) had two joinpoints – in 2008 and 2011. The values of ASRs increased by 17.7% per year from 2003 to 2008, decreased by 17.9% per year between 2008 and 2011, then increased by 17.0% per year from 2011 onwards. The ASRs at the national level (in Poland) showed an increasing pattern. The increased of 53.6% per year was observed between 1999 and 2001, 15.3% per year from 2001 to 2010, and 5.8% between 2010 and 2014.

Discussion

This is the most comprehensive descriptive study on breast cancer incidence in Poland that includes the examination of trends within voivodeships. The AAPCs indicate an overall increase in invasive and *in situ* breast cancer rates in Poland overall and within each of the voivodeships from 1999 to 2014. In Poland overall, incidence rates of invasive breast cancer in particular have significantly risen within the years 1999 to 2014, although the largest APCs have occurred prior to 2001. For voivodeships with joinpoints, the APC of invasive breast cancer incidence rates has consistently decreased, and for *in situ* breast cancer it has increased, probably reflecting the effects of breast cancer screening.

Incidence trends vary within each voivodeship. Since 2004 Łódź voivodeship has had the highest invasive breast cancer incidence rate, while Świętokrzyskie Voivodeship has had the highest *in situ* breast cancer incidence rate in the analysed periods of time. Łódź and Lublin voivodeships had the largest increase *in situ* and invasive breast cancer incidence rates.

An increase in breast cancer incidence rates was expected after implementation of the screening program in Poland. It was also observed in an earlier study by Botha *et al.* of 16 European countries that had adopted mammography screening in the 1980s [14]. The estimated APC ranges from 0.8% to 2.8% in

six 'screened' countries during pre-screening years, and from 1.2% to 3.0% in 10 'non-screened' countries [14]. Breast cancer incidence increased between 30% and 40% from the 1970s to the 1990s in most countries, with a visible increase among women aged 50 years and older. Breast cancer incidence rates are definitely higher in developed countries in comparison to developing countries, which is caused by distinct use of screening mammograms, discrepancies in lifestyle, and genetic factors [15].

With regards to breast cancer, Poland has historically had a high mortality-to-incidence ratio. The low incidence of breast cancer, in addition to the high mortality rate, low 5-year survival rates, and low morbidity are all an indication of ineffective primary cancer prevention campaigns, early detection, and treatment *per se* [16]. A similar situation also exists in other, recent members of the European Union, including Baltic countries, Bulgaria, and Romania. Among those countries, only Poland has implemented the National Programme of Cancer Prevention. One of the main goals of the National Health Programme in Poland was to increase early diagnosis and effectiveness of treatment of breast cancer [17]. The National Health Programme was beneficial in increasing the awareness of breast cancer prevention in Poland since its implementation [18]. From 1998 to 2001, the proportion of Polish women undergoing screening mammography nearly tripled from 10% to 28% [19]. One year after the accession of Poland into the European Union (EU), the Polish government enacted the bill implementing the National Programme Against Cancer Diseases (NPACD) for the years 2005–2010. In addition to increasing early detection of breast cancer, the goals of the NPACD included adopting diagnostic standards according to the EU [20].

On the other hand, mortality rates undergo gradual reductions. Increasing breast cancer incidence and a stable mortality trend shows the improvement of detection and treatment. During the last decade, an improvement of survival rates was observed in Poland, from 62% to 75%.

According to the observed trends in Poland and in selected voivodeships, breast cancer trends increased overall, although the trend was not consistent across all voivodeships. As the wealthiest and most urban voivodeship in Poland, Mazovia has the highest incidence rates of both invasive and *in situ* breast cancers. This is consistent with other studies, which have also found higher breast cancer rates in the urban population [21]. Conversely, the lowest incidence of breast cancer was found in Podlasie, Lublin, and Świętokrzyskie voivodeships among women aged 65+ years. The abovementioned regions are rural and during last 20 years there have been no large factories or industry.

Likewise, health benefits payer has presented results that indicate an increase in breast cancer 5-year

survival rates in the years 2005 to 2008, but there are huge differences in individual voivodeships – from 66% to 79% and from 75.3% to 82.4% [22, 23]. Significant differences in results, methods, and costs may show the usage of various procedures and financial models. Considering the relationship between therapeutic effects and bearing costs, women with breast cancer have no equal access to the health care system, depending on their place of residence.

Disparities among voivodeships may be the result of environmental and cultural causations. There are some reasons for the increasing health hazard for cancer diseases: improper nutrition (excessive animal fat intake and low vegetable and fruit intake), excessive alcohol intake, poor physical activity, exposure to carcinogens in the workplace, lack of awareness of cancer danger, and community aversion to screening programs [24]. There is a need to develop health education, smoking restrictions, and to create appropriate dietary habits and reduce harmful carcinogenic conditions in the workplace and residence. Differences in patterns probably reflect changes in lifestyle risk factors such as reproduction behaviours and diet but cannot exclude the possibility of higher screening mammography rates and early detection in some voivodeships [25].

Limitations of our study are similar to other studies using data based on registries. These include analysis of data only for women who have visited a doctor. Also, trend analysis is not sufficient to determine the cause of the increase. Relevant and established breast cancer risk factors, such as nulliparity, obesity, lack of exercise, young age at menopause, long-term use of hormone replacement therapy, personal family history, and others, could not have been included in this analysis. Thirdly, histological classifications of breast cancer have changed over the course of 14 years, which may result in some outcome misclassification.

This study also has several strengths. We covered 16 years of breast cancer incidence in Poland, which spans the course of political, economic, as well as cultural and lifestyle changes that have occurred. The use of a joinpoint regression analysis using NCR data allows us to obtain an objective representation of breast cancer incidence as well as a direct comparison across regions of varying geographical locations within Poland. Lastly, this is the only study to analyse cancer incidence rates across different voivodeships within new EU member countries and provides a broad insight as to the circumstances surrounding medium-level incidence rates in similar countries.

Conclusions

In situ and invasive breast cancer incidence rates in Poland increased between 1999 and 2014. The rise in invasive breast cancer incidence rates in Poland is largely due to improved screening as well as a shift in

lifestyle risk factors after the accession of Poland into the EU in 2004. The APCs have slowed since 2004 in each voivodeship but not in Poland overall. It remains uncertain if breast cancer incidence rates will continue to increase or stabilise over time. It will be very important to monitor breast cancer incidence so that resources can be put into place to identify voivodeships most in need of mammography screening and breast cancer awareness. Increased public awareness of the benefits of mammography screening, self-examination, and risk factors will be important in reducing breast cancer rate disparities between Poland and its neighbouring European Union states.

Conflict of interest

The authors declare no conflict of interest.

References

1. Yip CH, Buccimazza I, Hartman M, Deo SVS, Cheung PS. Improving outcomes in breast cancer for low and middle income countries. *World J Surg* 2015; 39: 686-692.
2. Beiki O, Hall P, Ekblom A, Moradi T. Breast cancer incidence and case fatality among 4.7 million women in relation to social and ethnic background: a population-based cohort study. *Breast Cancer Res* 2012; 14: R5.
3. Ferlay J, Soerjomataram I, Ervik M, Dikshit R, Eser S, Mathers C. GLOBOCAN 2012 v1. 0, cancer incidence and mortality worldwide: IARC CancerBase No. 11. International Agency for Research on Cancer, Lyon [online] [cited 2014 04.05]. Globocan Iarc Fr.
4. Jokiel M. Social aspects of breast cancer early detection after introduction of population screenings in Poland. *Przegl Epidemiol* 2008; 63: 443-447.
5. Liong YV, Hong GS, Teo JG, Lim GH. Breast ductal carcinoma in situ presenting as recurrent non-puerperal mastitis: case report and literature review. *World J Surg Oncol* 2013; 11: 179.
6. Survival Analysis 2000 – 2007 [online] [cited: 2015 10.30]. Available from: <https://w3.iss.it/site/EU5Results/forms/SA0007.aspx>.
7. Sytuacja demograficzna osób starszych i konsekwencje starzenia się ludności Polski w świetle prognozy na lata 2014-2050. Warszawa: Główny Urząd Statystyczny 2014; 1-43.
8. Demographic Yearbook of Poland 2014 [online] [cited: 2015 10.30]. Available from: <http://stat.gov.pl/en/topics/statistical-yearbooks/statistical-yearbooks/demographic-yearbook-of-poland-2014,3,8.html>
9. Raporty Krajowego Rejestru Nowotworów [online] [cited: 2015 03.20]. Available at: <http://onkologia.org.pl/raporty/>.
10. Didkowska J, Wojciechowska U, Zatoński W. Nowotwory złośliwe w Polsce w 2011 roku. *Cent Onkol Warszawa* 2013 [online] [cited: 15 marzec 2015]; Available at: http://onkologia.org.pl/wp-content/uploads/COI_Nowotwory2013_web.pdf.
11. Didkowska J, Wojciechowska U, Zatoński W. Prognozy zachorowalności i umieralności na nowotwory złośliwe w Polsce do 2025 roku. *Centrum Onkologii – Instytut im. M. Skłodowskiej-Curie*, Warsaw 2009.
12. Kim HJ, Fay MP, Feuer EJ, Midthune DN. Permutation tests for joinpoint regression with applications to cancer rates. *Stat Med* 2000; 19: 335-351.
13. Clegg LX, Hankey BF, Tiwari R, Feuer EJ, Edwards BK. Estimating average annual per cent change in trend analysis. *Stat Med* 2009; 28: 3670-3682.
14. Botha JL, Bray F, Sankila R, Parkin DM. Breast cancer incidence and mortality trends in 16 European countries. *Eur J Cancer* 2003; 39: 1718-1729.
15. Althuis MD, Dozier JM, Anderson WF, Devesa SS, Brinton LA. Global trends in breast cancer incidence and mortality 1973–1997. *Int J Epidemiol* 2005; 34: 405-412.
16. Curado MP. Breast cancer in the world: incidence and mortality. *Salud Pública México* 2011; 53: 372-384.
17. Department of Medical Statistics, National Institute of Hygiene. Health programme for years 1996–2005 [online] [cited: 2015 08.06]. Available at: http://www.medstat.waw.pl/nhp_p/1_p.html.
18. Hendrick RE, Smith RA, Rutledge 3rd JH, Smart CR. Benefit of screening mammography in women aged 40-49: a new meta-analysis of randomized controlled trials. *J Natl Cancer Inst Monogr* 1996; 22: 87-92.
19. Syczewska-Weber K, Rucinski P. The main challenges of Polish oncology. *Public Health Rep* 2008; 123: 655-63.
20. Act of 1 July 2005 on Establishing the Multi-Year “National Cancer Control Programme” [online] [cited 2015 08.06]. Available at: http://www.epaac.eu/from_heidi_wiki/Poland_Establishing_the_Multi-Year_National_Cancer_Control_Programme_English.pdf.
21. Krzyzak M, Maslach D, Juczewska M, Lasota W, Rabczenko D, Marcinkowski JT, Szpak A. Differences in breast cancer incidence and stage distribution between urban and rural female population in Podlaskie Voivodship, Poland in years 2001-2002. *Ann Agric Environ Med* 2010; 17: 159-162.
22. Kozierekiewicz A, Topór-Mądry R, Śliwczyński A, Pakulski M, Jassem J. Skuteczność i koszty leczenia raka piersi w Polsce: podejście regionalne. *Nowotw J Oncol* 2014; 64: 24-32.
23. Herman K, Śliwczyński A, Wysocki WM. Wyniki, metody i koszty leczenia raka piersi w Polsce (w latach 2005–2007). *Nowotw J Oncol* 2014; 64: 33-39.
24. Zatoński W, Przewoźniak K. Ograniczanie zachorowalności i umieralności z powodu chorób nowotworowych. [In:] *Zdrowie publiczne i polityka ludnościowa*. Rządowa Rada Ludnościowa. Szymborski J (ed.). Warsaw 2012; 78-89.
25. Ferlay J, Steliarova-Foucher E, Lortet-Tieulent J, Rosso S, Coebergh JW, Comber H, Forman D, Bray F. Cancer incidence and mortality patterns in Europe: estimates for 40 countries in 2012. *Eur J Cancer* 2013; 49: 1374-1403.

Address for correspondence:

Paweł Macek MD, PhD
 Department of Epidemiology and Cancer Control
 Holycross Cancer Centre
 ul. Artwińskiego 3, 25-734 Kielce, Poland
 Phone: +48 603 161 576
 E-mail: pawel.macek@gazeta.pl