

Incidence and survival of ocular melanoma in National Cancer Registry of Poland in 2010–2017

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Conflict of interest

None declared

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Abstract

Background. Oncology trends are based on data coming from different countries and ocular melanoma is the most common primary eye cancer in adults.

Objectives. To investigate the incidence and characteristics of ocular melanoma in the overall population of Poland.

Materials and methods. The retrospective survey of both the National Cancer Registry (NCR) and National Health Fund (NHF) databases was performed to identify all ocular melanoma cases in Poland in 2010–2017.

Results. The mean incidence of ocular melanoma was 8.76/1,000,000 person-years; the lowest incidence was observed in the 19–29 age group (1.17/1,000,000 person-years) and the highest in the group over 70 (22.88/1,000,000 person-years). There were no statistically significant trends in the incidence rates over the study period. The overall incidences of uveal, eyelid and conjunctival melanoma were 6.67/1,000,000, 0.47/1,000,000 and 0.28/1,000,000 person-years, respectively. The 5-year overall survival (OS) was 60.76%; the higher risk of death was associated with male sex (hazard ratio (HR) = 1.2959), older age at diagnosis (HR = 1.0379), chemotherapy treatment (HR = 1.6774), metastasis (HR = 1.5716), loco-regional hyperplasia (HR = 1.5936), and systemic tumor spread (HR = 3.9872), compared to the carcinoma in situ. The risk of death was reduced by radiotherapy treatment (HR = 0.6645).

Conclusions. The incidence rate of ocular melanoma in Poland is in the middle of the range worldwide, and the 5-year OS is relatively low.

Key words: radiotherapy, patient survival, ocular melanoma

Background

Ocular melanoma is the most common primary eye cancer in adults arising from melanocytes located in the conjunctival membrane and uveal tract of the eye, which accounts for 3.7% of all melanoma cases. Among ocular melanomas, 83% arise from the uvea, 5% from the conjunctiva and 10% from other sites in the eye. Uveal melanoma is the most common primary intraocular cancer in adults.^{1–5} The incidence of ocular melanoma varies across ethnicities and regions worldwide, with the highest rates in Northern Europe and Australia and the lowest rates in Asian, Hispanic and black populations (i.e., the incidence of uveal melanoma ranges from 0.31 in Black and 0.38 in Asian populations to 11.7/1,000,000 person-years in Northern Europe). However, the incidence rate of uveal melanoma remained stable over the recent decades, while conjunctival melanoma showed an increase in the incidence rate among white men and individuals over 60.^{1,4,6–10} The occurrence rate of ocular melanoma is positively correlated with older age, with a peak around the age of 70. However, the mean age of the ocular melanoma diagnosis also varies in different populations, i.e., in Asia, as it affects younger individuals than in Europe or the USA, where it usually presents around the age of 60. The anterior uveal melanoma is also more common in young patients. It represents more than 20% of all uveal melanomas in the age group under 20 compared to 4% and 2% in patients aged 20–60 and over 60, respectively.^{1,3,6} Other risk factors associated with ocular melanoma include sex, Caucasian origin, light skin and iris pigmentation, genetic predisposition (BAP1 mutation), environmental factors, and certain dermatological conditions like dysplastic nevus syndrome or nevus of Ota.^{11–14} Despite the fact that the treatment of ocular melanoma has evolved with a therapeutic shift to eye-conserving treatment options, it is estimated that still more than 50% of patients develop metastases within 25 years from the initial diagnosis. Poor prognosis indicators include older age at diagnosis, large tumor diameter, anterior location, extraocular extension, histopathological type, and cytogenetic abnormalities.^{3,8,15–19} Although the local data from Poland concerning conjunctival and uveal melanomas were analyzed by the RARECAREnet and the European Cancer Registry (EUROCARE) working groups,^{4,7,20} data from the entire population of Poland are lacking.

Objectives

The present study aims to analyze the incidence and characteristics of ocular melanoma in the overall population of Poland in 2010–2017, and to report the patient survival and coexisting risk factors.

Materials and methods

Data sources, disease codes and definitions

The present study was a part of the project “Maps of Healthcare Needs – Database of Systemic and Implementation Analyses” coordinated by the Polish Ministry of Health and co-financed by the European Union funds through the European Social Fund under the Knowledge, Education and Development Operational Program (EU grant No. POWR 05.02.00-00-0149/15-01).^{21–24} The study design was a retrospective and nationwide survey concerning patients with ocular melanoma diagnosed between January 1, 2010 and December 31, 2017. Patients were identified in the National Cancer Registry (NCR) using personal identification number (PESEL), the 10th revision of the International Statistical Classification of Diseases and Related Health Problems (ICD-10), and the 3rd edition of the International Classification of Diseases for Oncology (ICD-O-3) codes. The ICD-10 codes defining ocular neoplasms are C43.1 and C69 with extensions. Ocular melanomas were found in this group of patients using ICD-O-3 codes 872–877 with extensions. Therefore, all patients with both the ICD-10 and ICD-O-3 codes (as mentioned above) were included in the study group. The analysis of the study group also included demographic data from the National Health Fund (NHF) database, such as patient sex, age at diagnosis and area code. As both the NCR and the NHF databases cover the entire population of Poland, we believe that all ocular melanoma cases diagnosed in Poland between 2010 and 2017 were included in the statistical analysis. In addition, population data for Poland and patient death records were obtained from Statistics Poland.^{25–27}

Data analyses

In the 1st part of the study, the descriptive statistics and demographic characteristics of the study group and the incidence analysis of ocular melanoma were performed for each year of the study period, separately. Then, the analysis of patients diagnosed with ocular melanoma in 2010–2014 with 5 years of follow-up was carried out, allowing the performance of survival analysis. The 1-year and 5-year overall survival (OS) rates were calculated. The Cox proportional hazards model was applied, and hazard ratios (HRs) with a 95% confidence interval (95% CI) were computed. A value of $p < 0.05$ was considered statistically significant. The Kaplan–Meier curve was employed to present the 5-year survival. Many different factors were considered in the survival analysis, including patient sex, age at the time of diagnosis, place of residence, and treatment method. Data on the general treatment schedule were available in the NCR database; these data concern surgical treatment, radiotherapy, chemotherapy, and other therapies. Additionally, detailed therapies were identified using

the ICD-9 codes according to the ICD-9-CM Volume 3 classification (a subset of the International Statistical Classification of Diseases and Related Health Problems (ICD)-9-CM) and obtained from the NHF database of medical services. They were matched with the study group from NCR. The ICD-9 codes, namely 16.31, 16.39, 16.41, 16.42, 16.49, 16.51, and 16.52, were used to identify surgical treatment by enucleation, and ICD-9 codes 14.26, 14.27 and 92.4 with extensions were applied to identify radiotherapy by plaque brachytherapy. Other therapies indicated in the NCR database included laser therapy and targeted therapy. Other clinical factors obtained from the NCR database were also taken into account, including metastases or advancement stages. Tumor growth advancement was divided into 4 categories: in situ, local, loco-regional, and systemic. The reference group was the tumor in situ. The R statistical software v. 3.6.2 (R Foundation for Statistical Computing, Vienna, Austria) was used for all analyses. The demographic characteristics of patients are presented with the mean and standard deviation (SD).

Since our study did not require ethics committee approval, it adhered to the tenets of the Declaration of Helsinki for research involving human subjects (socio-demographic data, including age, sex and place of residence, were recorded anonymously). Furthermore, the study protocol was approved by the Polish Ministry of Health, which is entitled by the laws of the Republic of Poland to process the NHF data. The informed consent was waived.

Results

In total, 2143 patients with ocular melanoma were identified in Poland between January 1, 2010 and December 31, 2017. The incidence rates for each age group in the study period are presented in Table 1 and Fig. 1. The mean incidence of ocular melanoma was 8.76/1,000,000 person-years (95% CI: [6.94; 10.58]). The incidence of ocular melanoma increased with age: the lowest incidence was observed in patients aged 19–29 (1.17/1,000,000 person-years) and the highest in patients over

Table 1. Age-standardized incidence of ocular melanoma among Polish adults from 2010 to 2017 by age group

Variable	2010	2011	2012	2013	2014	2015	2016	2017	All
Number of people aged 19–29 years (in thousands)	6117.1	6015.0	5854.2	5651.8	5451.4	5243.7	5062.3	4890.1	44,285.7
Number of melanoma cases	9	11	8	4	6	6	2	6	52
Incidence/1,000,000 person-years	1.47	1.83	1.37	0.71	1.10	1.14	0.39	1.23	1.17
Percentage of women [%]	33.33	45.45	50.00	25.00	50.00	83.33	0.00	83.33	50.00
Number of people aged 30–39 years (in thousands)	5895.1	6005.6	6123.5	6239.5	6314.5	6348.3	6330.6	6290.1	49,547.1
Number of melanoma cases	15	12	14	12	14	15	15	12	109
Incidence/1,000,000 person-years	2.54	2	2.29	1.92	2.22	2.36	2.37	1.91	2.2
Percentage of women [%]	40.00	66.67	64.29	83.33	28.57	46.67	60.00	41.67	53.21
Number of people aged 40–49 years (in thousands)	4847.2	4822.2	4838.4	4879.8	4956.0	5064.6	5202.4	5341.5	39,952.2
Number of melanoma cases	28	29	33	10	17	19	22	37	195
Incidence/1,000,000 person-years	5.78	6.01	6.82	2.05	3.43	3.75	4.23	6.93	4.88
Percentage of women [%]	46.43	68.97	48.48	70.00	76.47	57.89	40.91	43.24	53.85
Number of people aged 50–59 years (in thousands)	5847.7	5765.5	5656.6	5536.1	5406.3	5245.3	5089.3	4928.3	43,475.2
Number of melanoma cases	63	47	49	44	42	50	61	81	437
Incidence/1,000,000 person-years	10.77	8.15	8.66	7.95	7.77	9.53	11.99	16.44	10.05
Percentage of women [%]	58.73	55.32	44.90	50.00	59.52	52.00	55.74	58.02	54.69
Number of people aged 60–69 years (in thousands)	3712.0	3,931.3	4,171.2	4,409.8	4,642.8	4,888.3	5,024.7	5,127.3	35,907.4
Number of melanoma cases	68	82	77	51	53	96	88	115	630
Incidence/1,000,000 person-years	18.32	20.86	18.46	11.57	11.42	19.64	17.51	22.43	17.55
Percentage of women [%]	41.18	40.24	57.14	54.90	41.51	57.29	45.45	51.30	49.05
Number of people aged ≥70 years (in thousands)	3830.6	3865.3	3874.0	3882.9	3905.0	3914.7	4030.5	4166.8	31,469.2
Number of melanoma cases	78	80	94	68	74	85	101	140	720
Incidence/1,000,000 person-years	20.36	20.7	24.26	17.51	18.95	21.71	25.06	33.6	22.88
Percentage of women [%]	53.26	54.41	51.27	58.20	52.91	55.35	52.60	52.43	57.08
Total number of people in all above age groups combined (in thousands)	30,249.8	30,404.7	30,518.0	30,599.9	30,675.9	30,705.0	30,739.9	30,743.6	244,636.8
Number of melanoma cases	261	261	275	189	206	271	289	391	2143
Incidence/1,000,000 person-years	8.63	8.58	9.01	6.18	6.71	8.83	9.40	12.72	8.76
Percentage of women [%]	53.26	54.41	51.27	58.20	52.91	55.35	52.60	52.43	53.57

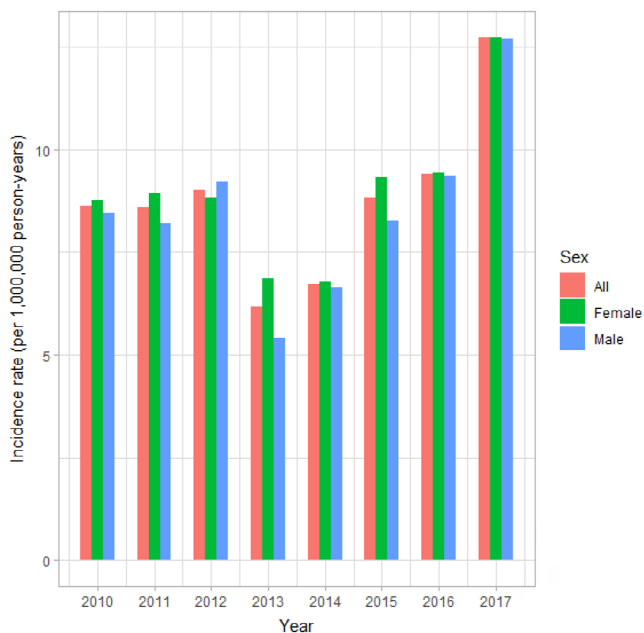


Fig. 1. Incidence of ocular melanoma in Poland according to standard annual analysis during 2010–2017

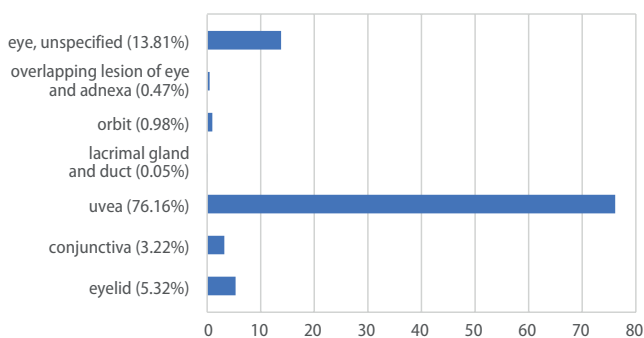


Fig. 2. Localization of ocular melanoma among Polish adults from 2010 to 2017

70 (22.88/1,000,000 person-years). The number of ocular melanoma diagnoses by anatomical localization is presented in Fig. 2. The uveal melanoma (iris, ciliary body and choroid localizations) was the most common and was diagnosed in 1632 (76.16%) patients. Eyelid melanoma was the 2nd most common and was diagnosed in 114 (5.32%) patients. The 3rd

most common was conjunctival melanoma, which was diagnosed in 69 (3.22%) patients. The other melanomas were localized in orbit (21 patients (0.98%)), overlapping lesion of eye and adnexa (10 patients (0.47%)) and lacrimal gland and duct (1 patient (0.05%)). At the same time, unspecified ocular melanoma was diagnosed in 296 (13.81%) patients. The overall incidences of uveal, eyelid and conjunctival melanoma were 6.67/1,000,000, 0.47/1,000,000 and 0.28/1,000,000 person-years, respectively.

The demographic characteristics of patients with ocular melanoma in Poland are presented in Fig. 1 and Table 1,2. The mean age at the time of diagnosis was 62.73 ± 14.43 years, and there was a slight increase in the mean age of diagnosis over the study period. The majority of patients were female (53.57%), and the proportion of women in each analyzed year was more than half. In addition, the vast majority of patients (65%) were urban residents. The overall ocular melanoma incidence rate was also higher in women (8.96/1,000,000 person-years) than in men (8.54/1,000,000 person-years).

The OS analysis included 1192 patients diagnosed in 2010–2014. One hundred patients (8.39%) died within 1 year and 507 patients (39.24%) died within 5 years from ocular melanoma diagnosis. The 1-year and 5-year OS rates were 91.61% and 60.76%, respectively. Among statistically significant variables, a higher risk of death within 5 years was associated with male sex (HR = 1.2959; 95% CI: [1.086; 1.547]), older age at diagnosis (HR = 1.0379; 95% CI: [1.03; 1.046]), chemotherapy treatment (HR = 1.6774; 95% CI: [1.317; 2.136]), metastasis (HR = 1.5716; 95% CI: [1.056; 2.340]), loco-regional hyperplasia (HR = 1.5936; 95% CI: [1.111; 2.286]), and systemic tumor spread (HR = 3.9872; 95% CI: [3.021; 5.263]) compared to the carcinoma in situ. The risk of death was statistically reduced by radiotherapy treatment (HR = 0.6645; 95% CI: [0.529; 0.835]). However, place of residence, surgery treatment and local tumor growth (compared to the tumor in situ) were not associated with the mortality rate. All results of survival analysis are presented in Table 3. The Kaplan–Meier curve shows the patient survival (Fig. 3). The results of the test for the proportional hazards assumption are presented in Table 4.

Table 2. Demographic characteristics of patients with ocular melanoma among Polish adults from 2010 to 2017

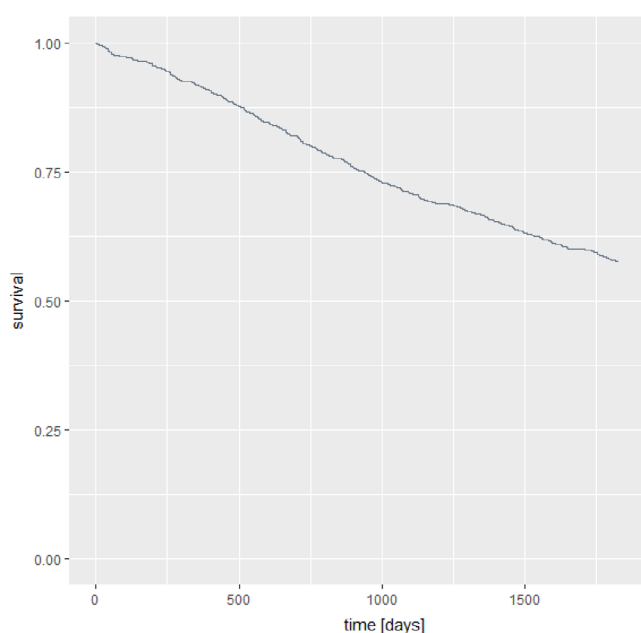
Variable	2010	2011	2012	2013	2014	2015	2016	2017	All
Age [years], mean \pm SE	60.72 \pm 14.88	60.68 \pm 14.83	62.17 \pm 14.39	63.47 \pm 14.26	62.59 \pm 15.53	63.46 \pm 14.09	63.92 \pm 14.10	64.18 \pm 13.61	62.73 \pm 14.43
Women, n (%)	139 (53.26)	142 (54.41)	141 (51.27)	110 (58.20)	109 (52.91)	150 (55.35)	152 (52.60)	205 (52.43)	1148 (53.57)
Men, n (%)	122 (46.74)	119 (45.59)	134 (48.73)	79 (41.80)	97 (47.09)	121 (44.65)	137 (47.40)	186 (47.57)	995 (46.43)
Urban residence, n (%)	163 (62.45)	181 (69.35)	167 (60.73)	122 (64.55)	121 (58.74)	175 (64.58)	196 (67.82)	268 (68.54)	1393 (65.00)
Rural residence, n (%)	98 (37.55)	80 (30.65)	108 (39.27)	67 (35.45)	85 (41.26)	96 (35.42)	93 (32.18)	123 (31.46)	750 (35.00)

SE – standard error.

Table 3. Results of Cox proportional hazards model of 5-year survival

Variable	B	SE	p-value	HR	2.5% HR	97.5% HR
Rural residence	−0.012038	0.092972	0.896974	0.9880	0.8234	1.1855
Male sex	0.259177	0.090260	0.004086	1.2959	1.0857	1.5466
Age	0.037241	0.003708	<0.00001	1.0379	1.0304	1.0455
Surgery	0.139545	0.111631	0.211279	1.1498	0.9238	1.4310
Chemotherapy	0.517221	0.123385	0.0000277	1.6774	1.3170	2.1362
Radiotherapy	−0.408784	0.116795	0.000465	0.6645	0.5285	0.8354
Metastasis	0.452075	0.203065	0.025997	1.5716	1.0556	2.3398
Local cancer	−0.107813	0.110736	0.330253	0.8978	0.7226	1.1154
Loco-regional cancer	0.465971	0.184095	0.011369	1.5936	1.1109	2.2860
Systemic cancer	1.383081	0.141593	<0.00001	3.9872	3.0209	5.2625

SE – standard error; HR – hazard ratio.

**Fig. 3.** Kaplan–Meier curve of ocular melanoma survival in Poland

Discussion

This study evaluates for the first time the incidence and characteristics of ocular melanoma in the overall population of Poland. The analysis included both the age-standardized incidence of ocular melanoma (the unit of incidence is captured by 1,000,000 person-years) in 2010–2017 and the survival analysis of patients diagnosed with ocular melanoma in 2010–2014, with 5 years of follow-up. The mean incidence of ocular melanoma was 8.76/1,000,000 person-years, and the mean age at the time of diagnosis was 62.73 ± 14.43 years. The overall incidences of uveal, eyelid and conjunctival melanoma were 6.67/1,000,000, 0.47/1,000,000 and 0.28/1,000,000 person-years, respectively. Our results were in the middle of the range of ocular melanoma incidence worldwide, which is much higher among Whites than Blacks and Asians.^{1–3} Our age-standardized incidence of uveal melanoma was similar to that reported earlier

in Poland by the EURO CARE working group for the 1983–1994 period.⁷ It was also similar to that reported in Central Europe (Slovakia, Slovenia and Switzerland), higher than that reported in South Korea, Singapore, USA, and Southern Europe (Italy and Spain) but lower than in Northern Europe (Denmark, Norway, Sweden, and Estonia), Ireland and Australia.^{6,7,9,11,15} However, we cannot exclude the misclassification bias. The number of uveal melanoma cases might have been underestimated, while unspecified ocular melanoma might have been overestimated. Errors in using specific ICD-10 codes might have occurred at different levels (hospitals, outpatient clinics, and NCR and NHF offices). However, we believe that such errors had only minor impact on the study findings. Our age-standardized incidence of conjunctival melanoma was similar to that reported earlier in Eastern Europe by the EURO CAREnet working group for the 1995–2007 period and higher than that reported among Blacks in the USA and whole population in Southern Europe but lower than among Whites in the USA and the whole population in Northern Europe.⁴ The north-to-south decreasing gradient in the uveal melanoma incidence in Europe might be related to the protective effect of ocular pigmentation in the southern populations with respect to higher exposure to ultraviolet light at lower latitudes. Moreover, Eastern European countries like Poland have the lowest rates of both conjunctival and skin melanomas in Europe.^{4,7} Most of the previous studies showed an increased age-adjusted incidence rate of ocular melanoma among men,^{6,9,11,12,14,15,17} but in other large cohort clinical studies with no age adjustment, no sex-based differences were reported.³ In contrast to those studies, the incidence rate of ocular melanoma in Poland was higher in women, which might be attributable to the excess male death rate, characteristic to the Eastern European countries, which is visible in Poland.^{27–29}

Since the Collaborative Ocular Melanoma Study (COMS) showed no survival advantage of enucleation over brachytherapy for medium-size tumors, the therapeutic shift to eye-conserving treatment options was observed worldwide.^{17,19} However, medical management of ocular

Table 4. Results of the test for the proportional hazards assumption

Variable	χ^2	Degree of freedom (df)	p-value
Rural residence	0.0754	1	0.7837
Male sex	1.2156	1	0.2702
Age	0.3953	1	0.5295
Surgery	1.182	1	0.277
Chemotherapy	1.9842	1	0.1590
Radiotherapy	0.2413	1	0.6233
Metastasis	1.3794	1	0.2402
Local cancer	0.3865	1	0.534
Loco-regional cancer	0.2574	1	0.6119
Systemic cancer	1.9519	1	0.162
Global	11.80962	10	0.212

melanoma depends on the tumor location, size, local extension, visual acuity at presentation, and systemic status. Most patients with posterior tumors are currently treated with plaque brachytherapy. Other available options include laser photocoagulation, transpupillary thermotherapy, particle beam radiotherapy, gamma knife radiosurgery, local surgical resection, and/or enucleation (with or without orbital exenteration). The current standard for anterior tumors management is a surgical treatment with adjuvant therapy, including brachytherapy. Other management methods for both anterior and posterior tumors include targeted therapy and/or chemotherapy (in case of metastases). Although more than 90% of primary tumors are managed with surgery or eye-conserving therapies, more than 50% of patients develop metastases, usually involving the liver.^{1,3,16,30}

The OS analysis revealed that 100 patients (8.39%) died within 1 year and 507 patients (39.24%) died within 5 years from the initial diagnosis of ocular melanoma in Poland, which gives the 1-year and 5-year mortality rates of 8.39% and 39.24%, respectively. The 1-year OS was 91.61%, and the 5-year OS was 60.76%, which is comparable to the data from an epidemiological study of uveal melanoma from US Surveillance, Epidemiology and End Results Program for 2010–2015, where the 5-year OS was 61.8%. However, our mortality rate was higher than that found in the UK, Denmark, Sweden, Singapore, or Israel.^{2,10,14,20,31,32} The Cox proportional hazards model showed that the higher risk of death within 5 years from the initial ocular melanoma diagnosis in Poland was associated with male sex, older age at diagnosis, chemotherapy treatment, metastasis, loco-regional hyperplasia, and systemic tumor spread compared to the carcinoma in situ. The risk of death was statistically reduced by radiotherapy treatment. However, chemotherapy treatment increased the risk of death in our model. It should be assumed that the treatment method depends on the carcinoma stage and chemotherapy is used in more severe stage of tumor. Our results were consistent with previously published studies that showed older age

at diagnosis, severe stage of the tumor, distant metastasis, and no radiation to be associated with the mortality risk. Those studies also revealed that beyond 15 years from diagnosis, a patient with uveal melanoma is more likely to die from other causes than uveal melanoma metastasis itself.^{14,31,32}

Limitations

The limitations of the present study include selection bias as there is an increasing proportion of cases without histopathological proof of diagnosis. In addition, both NHF and NCR databases do not cover the family history and genetic information of ocular melanoma patients in Poland, with disease laterality also not available for all included subjects. Therefore, we might have missed some data and the investigation of potential risk factors of mortality was not complex. However, this likely had only a minor impact on our findings. The most important strengths of the present study are the population size, nationwide recruitment, and the usefulness of its results for clinicians and health-care providers in Poland.

Conclusions

This study found that the incidence rate of ocular melanoma in Poland is in the middle range of worldwide incidence, and the 5-year OS is relatively low. Furthermore, a higher risk of mortality from ocular melanoma in Poland is associated with male sex, older age at diagnosis, chemotherapy treatment, metastasis, loco-regional hyperplasia, and systemic tumor spread compared to carcinoma in situ.

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References

- Jovanovic P, Mihajlovic M, Djordjevic-Jocic J, Vljakovic S, Cekic S, Stefanovic V. Ocular melanoma: An overview of the current status. *Int J Clin Exp Pathol*. 2013;6(7):1230–1244. PMID:23826405, PMCID: PMC3693189.
- Tan LLY, Hong J, Goh WL, et al. Clinical features and survival outcomes of ocular melanoma in a multi-ethnic Asian cohort. *Sci Rep*. 2020;10(1):16367. doi:10.1038/s41598-020-73534-x
- Kaliki S, Shields CL. Uveal melanoma: Relatively rare but deadly cancer. *Eye (Lond)*. 2017;31(2):241–257. doi:10.1038/eye.2016.275
- Virgili G, Parravano M, Gatta G, et al; RARECAREnet Working Group. Incidence and survival of patients with conjunctival melanoma in Europe. *JAMA Ophthalmol*. 2020;138(6):601–608. doi:10.1001/jamaophthalmol.2020.0531
- Mierzwa-Dobranowska M, Romanowska-Dixon B. The impact of selected factors on early diagnosis of multiple primary cancers in patients with uveal melanoma. *Contemp Oncol (Pozn)*. 2013;17(6):510–514. doi:10.5114/wo.2013.38914
- Ortega MA, Fraile-Martínez O, García-Honduvilla N, et al. Update on uveal melanoma: Translational research from biology to clinical practice (review). *Int J Oncol*. 2020;57(6):1262–1279. doi:10.3892/ijo.2020.5140

7. Virgili G, Gatta G, Ciccolallo L, et al; EURO CARE Working Group. Incidence of uveal melanoma in Europe. *Ophthalmology*. 2007;114(12):2309–2315. doi:10.1016/j.ophtha.2007.01.032
8. Burr JM, Mitry E, Rachet B, Coleman MP. Survival from uveal melanoma in England and Wales 1986 to 2001. *Ophthalmic Epidemiol*. 2007;14(1):3–8. doi:10.1080/09286580600977281
9. Park SJ, Oh CM, Kim BW, Woo SJ, Cho H, Park KH. Nationwide incidence of ocular melanoma in South Korea by using the National Cancer Registry Database (1999–2011). *Invest Ophthalmol Vis Sci*. 2015;56(8):4719–4724. doi:10.1167/iovs.15-16532
10. Frenkel S, Hendler K, Peer J. Uveal melanoma in Israel in the last two decades: Characterization, treatment and prognosis. *Isr Med Assoc J*. 2009;11(5):280–285. PMID:19637505.
11. Stang A, Parkin DM, Ferlay J, Jöckel KH. International uveal melanoma incidence trends in view of a decreasing proportion of morphological verification. *Int J Cancer*. 2005;114(1):114–123. doi:10.1002/ijc.20690
12. Nichols EE, Richmond A, Daniels AB. Disparities in uveal melanoma: Patient characteristics. *Semin Ophthalmol*. 2016;31(4):296–303. doi:10.3109/08820538.2016.1154176
13. Lucena E, Goldemberg DC, Thuler LCS, de Melo AC. Epidemiology of uveal melanoma in Brazil. *Int J Retina Vitreous*. 2020;6(1):51. doi:10.1186/s40942-020-00261-w
14. Xu Y, Lou L, Wang Y, et al. Epidemiological study of uveal melanoma from US Surveillance, Epidemiology, and End Results Program (2010–2015). *J Ophthalmol*. 2020;2020:3614039. doi:10.1155/2020/3614039
15. Baily C, O'Neill V, Dunne M, et al. Uveal melanoma in Ireland. *Ocul Oncol Pathol*. 2019;5(3):195–204. doi:10.1159/000492391
16. Nichols EE, Richmond A, Daniels AB. Tumor characteristics, genetics, management, and the risk of metastasis in uveal melanoma. *Semin Ophthalmol*. 2016;31(4):304–309. doi:10.3109/08820538.2016.1154175
17. Aronow ME, Topham AK, Singh AD. Uveal melanoma: 5-year update on incidence, treatment, and survival (SEER 1973–2013). *Ocul Oncol Pathol*. 2018;4(3):145–151. doi:10.1159/000480640
18. Mahendraraj K, Lau CS, Lee I, Chamberlain RS. Trends in incidence, survival, and management of uveal melanoma: A population-based study of 7516 patients from the Surveillance, Epidemiology, and End Results database (1973–2012). *Clin Ophthalmol*. 2016;10:2113–2119. doi:10.2147/OPHTH.S113623
19. Scheffler AC, Kim RS. Recent advancements in the management of retinoblastoma and uveal melanoma. *Fac Rev*. 2021;10:51. doi:10.12703/r/10-51
20. Virgili G, Gatta G, Ciccolallo L, et al; EURO CARE Working Group. Survival in patients with uveal melanoma in Europe. *Arch Ophthalmol*. 2008;126(10):1413–1418. doi:10.1001/archophth.126.10.1413
21. Nowak MS, Grabska-Liberek I, Michalska-Małecka K, et al. Incidence and characteristics of cataract surgery in Poland during 2010–2015. *Int J Environ Res Public Health*. 2018;15(3):435. doi:10.3390/ijerph15030435
22. Nowak MS, Grzybowski A, Michalska-Małecka K, et al. Incidence and characteristics of endophthalmitis after cataract surgery in Poland during 2010–2015. *Int J Environ Res Public Health*. 2019;16(12):2188. doi:10.3390/ijerph16122188
23. Koziół M, Nowak MS, Udziela M, Piątkiewicz P, Grabska-Liberek I, Szaflik JP. First nation-wide study of diabetic retinopathy in Poland in the years 2013–2017. *Acta Diabetol*. 2020;57(10):1255–1264. doi:10.1007/s00592-020-01540-6
24. Nowak MS, Romanowska-Dixon B, Grabska-Liberek I, Żurek M. Incidence and characteristics of retinoblastoma in Poland: The first nationwide study 2010–2017. *Int J Environ Res Public Health*. 2021;18(12):6539. doi:10.3390/ijerph18126539
25. The National Cancer Registry Data. <http://onkologia.org.pl>. Accessed July 16, 2021.
26. The National Health Fund Data. <http://www.nfz.gov.pl>. Accessed July 16, 2021.
27. Statistics Poland Data. <http://www.stat.gov.pl>. Accessed July 16, 2021.
28. Nowak MS, Śmigiełski J. The prevalence and causes of visual impairment and blindness among older adults in the city of Lodz, Poland. *Medicine (Baltimore)*. 2015;94(5):e505. Erratum in: *Medicine (Baltimore)*. 2015;94(7):1. doi:10.1097/MD.0000000000000505
29. Nowak MS, Jurowski P, Gos R, Śmigiełski J. Ocular findings among young men: A 12-year prevalence study of military service in Poland. *Acta Ophthalmol*. 2010;88(5):535–540. doi:10.1111/j.1755-3768.2008.01476.x
30. Chattopadhyay C, Kim DW, Gombos DS, et al. Uveal melanoma: From diagnosis to treatment and the science in between. *Cancer*. 2016;122(15):2299–2312. doi:10.1002/cncr.29727
31. Rajeshuni N, Zubair T, Ludwig CA, Moshfeghi DM, Mruthyunjaya P. Evaluation of racial, ethnic, and socioeconomic associations with treatment and survival in uveal melanoma, 2004–2014. *JAMA Ophthalmol*. 2020;138(8):876–884. doi:10.1001/jamaophthalmol.2020.2254
32. Radivoyevitch T, Zabor EC, Singh AD. Uveal melanoma: Long-term survival. *PLoS One*. 2021;16(5):e0250939. doi:10.1371/journal.pone.0250939