

The Impact of Oophorectomy on Survival from Breast Cancer in Patients with *CHEK2* Mutations

Joanna Tomiczek-Szwiec,¹Marek Szwiec,^{2,3}Michał Falco,⁴Cezary Cybulski,⁵Dominika Wokolorczyk,⁵Anna Jakubowska,⁵Jacek Gronwald,⁵Małgorzata Stawicka,⁶Dariusz Godlewski,⁷Ewa Kilar,⁸Elżbieta Marczyk,⁹Monika Siołek,¹⁰Rafał Wiśniowski,¹¹Olga Haus,¹²Robert Sibilski,^{2,13}Lubomir Bodnar,¹⁴Ping Sun,¹⁵Steven A. Narod,¹⁵Jan Lubinski,⁵Tomasz Huzarski,^{5,6} and the Polish Breast Cancer Consortium*

¹Department of Histology, Department of Biology and Genetics, Faculty of Medicine, University of Opole, Opole, Poland

²Department of Surgery and Oncology, University of Zielona Góra, Zyty 28 St, 65-046 Zielona Góra, Poland

³Department of Clinical Oncology, University Hospital in Zielona Góra, Zyty 26 St, 65-046 Zielona Góra, Poland

⁴Regional Oncology Hospital, Strzalowska 22, 71-730 Szczecin, Poland

⁵Department of Genetics and Pathology, International Hereditary Cancer Center, Pomeranian Medical University, Unii Lubelskiej 1 St, 71-252 Szczecin, Poland

⁶Department of Clinical Genetics and Pathology, University of Zielona Góra, Zyty 28 St, 65-046 Zielona Góra, Poland

⁷OPEN, Kazimierza Wielkiego 24 St, 61-863 Poznań, Poland

⁸Department of Oncology, District Specialist Hospital, Leśna 27-29 St, 58-100 Świdnica, Poland

⁹Department of Oncological Surgery, Regional Oncology Center, Gancarska 11 St, 31-115 Kraków, Poland

¹⁰Holycross Cancer Center, Artwińskiego 3 St, 25-734 Kielce, Poland

¹¹Regional Oncology Hospital, Wyzwolenia 18 St, 43-300 Bielsko Biała, Poland

¹²Department of Clinical Genetics, Collegium Medicum, Nicolaus Copernicus University, Jagiellońska 13 St, 85-067 Bydgoszcz, Poland

¹³Oncology Diagnostic Center, Wazów 42 St, 65-044 Zielona Góra, Poland

¹⁴Department of Oncology and Immuno-oncology, School of Medicine, Collegium Medicum, University of Warmia and Mazury in Olsztyn, Warszawska 30 St, 10-082 Olsztyn, Poland

¹⁵Women's College Research Institute, Toronto, Ontario, Canada, M5G 1N8

Background: To estimate the impact of oophorectomy and other treatments on the survival of breast cancer patients with a *CHEK2* mutation.

Methods: Women with stage I-III breast cancer who were treated at 17 hospitals in Poland were tested for four founder mutations in the *CHEK2* gene. 974 women (10%) were positive for a *CHEK2* mutation. Control patients without a *CHEK2* mutation were selected from a database of patients treated over the same time period. Information on treatments received and distant recurrences was retrieved from medical records. Treatments included chemotherapy, hormonal therapy (tamoxifen) and radiation therapy. Oophorectomies were performed for the treatment of breast cancer or for benign conditions. Dates of death were obtained from the Polish Vital Statistics Registry. Causes of death were determined by medical record review. Predictors of survival were determined using the Cox proportional hazards model.

Results: 839 patients with a *CHEK2* mutation were matched to 839 patients without a mutation. The mean follow-up was 12.0 years. The 15-year survival for *CHEK2* carriers was 76.6% and the 15-year survival for non-carrier control patients was 78.8% (adjusted HR = 1.06; 95% CI 0.84 to 1.34; p = 0.61). Among *CHEK2* carriers, the 15-year survival for woman who had an oophorectomy was 86.3% and for women who did not have an oophorectomy was 72.1% (adjusted HR = 0.59; 95% CI 0.38 to 0.90; p = 0.02). Among controls, the 15-year survival for patients who had an oophorectomy was 84.5% and for women who did not have an oophorectomy was 77.6% (adjusted HR = 1.03; 95% CI 0.66 to 1.61; p = 0.90).

Conclusion: Among women with breast cancer and a *CHEK2* mutation, oophorectomy is associated with a reduced risk of death from breast cancer.