

Myungshin Kim, M.D., Ph.D.<sup>1,2</sup><sup>1</sup>Department of Laboratory Medicine, College of Medicine, The Catholic University of Korea, Seoul, Korea; <sup>2</sup>Catholic Genetic Laboratory Center, Seoul St. Mary's Hospital, College of Medicine, The Catholic University of Korea, Seoul, Korea

## Cost-effective *BRCA* Testing in Advanced Ovarian Cancer

Ovarian cancer is the most aggressive gynecological cancer worldwide [1]. Because of the deficiency in early detection procedures and rapid progression of the disease, more than 70% of ovarian cancer patients are only diagnosed at an advanced stage. In Korea, the incidence of ovarian cancer has gradually increased, partly due to the westernization of lifestyle and changes in reproductive factors, including early menarche, late menopause, delay in marriage, lower fertility rate in life, and changes in infant feeding patterns [2, 3]. Ovarian cancer is the most common cause of gynecological cancer-related deaths in Korea [4]. Despite advances in treatment over the past few decades, the 5-year survival rate remains below 40% [5]. Most recently, targeted agents, such as poly (ADP-ribose) polymerase (PARP) inhibitors, have driven improvement. PARP inhibitors were the first approved cancer drugs to specifically target the DNA damage response in *BRCA1/2*-mutated breast and ovarian cancers [6]. Since their introduction, the significance of *BRCA* mutations, not only for identifying hereditary breast and ovarian cancer syndromes but also for selecting patients suitable for targeted cancer therapy, has been greatly emphasized [7, 8].

Two types of samples can be used for *BRCA* testing: germline and somatic samples. Germline *BRCA* mutations are analyzed in peripheral blood samples, whereas somatic mutations are detected in tumor samples. In comparison with the analysis of DNA isolated from blood samples, testing DNA isolated from formalin-fixed paraffin-embedded (FFPE) tissue is challenging

because various factors, such as FFPE age, DNA fragmentation, and deamination, can reduce specimen integrity [9, 10]. Although *BRCA* testing using both samples is beneficial for ovarian cancer patients, because of the limited resources, it is desirable to develop an optimal *BRCA* testing strategy based on economic evaluation.

This issue was addressed in a recent study by Jang, *et al.* [11]. They assessed the cost-effectiveness of *BRCA* testing strategies followed by PARP inhibitor maintenance therapy based on the National Health Insurance system in Korea. They evaluated five *BRCA* testing strategies: (1) germline testing first, followed by somatic tumor testing for patients without germline mutations; (2) somatic testing first, followed by germline testing for patients with mutations detected by somatic testing; (3) both germline and somatic testing; (4) germline testing alone; and (5) somatic testing alone. By analyzing the incremental cost-effectiveness ratio, the authors found that all five strategies were cost-effective. Strategy 4 was the most cost-effective option, with an incremental cost-effectiveness ratio, followed by strategy 1. They concluded that germline testing first, followed by somatic testing, may be a reasonable *BRCA* testing option for Korean patients with advanced ovarian cancer. This study provides important and valuable information for clinicians, clinical laboratories, and the government to consider and plan *BRCA* testing strategies for advanced ovarian cancer in the targeted cancer therapy era.



© Korean Society for Laboratory Medicine

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (<https://creativecommons.org/licenses/by-nc/4.0>) which permits unrestricted non-commercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

## AUTHOR CONTRIBUTIONS

Kim M accepts responsibility for the content of this manuscript and approved its submission.

## CONFLICTS OF INTEREST

No potential conflicts of interest relevant to this paper were reported.

## ORCID

Myungshin Kim <https://orcid.org/0000-0001-8632-0168>

## REFERENCES

1. Bray F, Ferlay J, Soerjomataram I, Siegel RL, Torre LA, Jemal A. Global cancer statistics 2018: GLOBOCAN estimates of incidence and mortality worldwide for 36 cancers in 185 countries. *CA Cancer J Clin* 2018; 68:394-424.
2. Park B, Park S, Kim TJ, Ma SH, Kim BG, Kim YM, et al. Epidemiological characteristics of ovarian cancer in Korea. *J Gynecol Oncol* 2010;21: 241-7.
3. Hunn J and Rodriguez GC. Ovarian cancer: etiology, risk factors, and epidemiology. *Clin Obstet Gynecol* 2012;55:3-23.
4. Jung KW, Won YJ, Kong HJ, Lee ES. Prediction of cancer incidence and mortality in Korea, 2019. *Cancer Res Treat* 2019;51:431-7.
5. Wright JD, Chen L, Tergas AI, Patankar S, Burke WM, Hou JY, et al. Trends in relative survival for ovarian cancer from 1975 to 2011. *Obstet Gynecol* 2015;125:1345-52.
6. George A, Kaye S, Banerjee S. Delivering widespread *BRCA* testing and PARP inhibition to patients with ovarian cancer. *Nat Rev Clin Oncol* 2017; 14:284-96.
7. Colombo N, Huang G, Scambia G, Chalas E, Pignata S, Fiorica J, et al. Evaluation of a streamlined oncologist-led *BRCA* mutation testing and counseling model for patients with ovarian cancer. *J Clin Oncol* 2018; 36:1300-7.
8. Lee A, Moon BI, Kim TH. *BRCA1/BRCA2* pathogenic variant breast cancer: treatment and prevention strategies. *Ann Lab Med* 2020;40:114-21.
9. Lee A, Kang J, Lee H, Lee YS, Choi YJ, Lee KH, et al. *BRCA1/2* somatic mutation detection in formalin-fixed paraffin embedded tissue by next-generation sequencing in Korean ovarian cancer patients. *Pathol Res Pract* 2019;215:152595.
10. Yoo J, Lee GD, Kim JH, Lee SN, Chae H, Han E, et al. Clinical validity of next-generation sequencing multi-gene panel testing for detecting pathogenic variants in patients with hereditary breast-ovarian cancer syndrome. *Ann Lab Med* 2020;40:148-54.
11. Jang J, Kim Y, Kim JH, Cho SM, Lee KA. Cost-Effectiveness Analysis of Germline and Somatic *BRCA* Testing in Patients with Advanced Ovarian Cancer. *Ann Lab Med* 2023;43:73-81.

---

**Corresponding author: Myungshin Kim, M.D., Ph.D.**

Department of Laboratory Medicine, College of Medicine, The Catholic University of Korea, 222 Banpo-daero, Seocho-gu, Seoul 06591, Korea  
Tel: +82-2-2258-1645, Fax: +82-2-2258-1719  
E-mail: microkim@catholic.ac.kr

**Key Words:** *BRCA1*, *BRCA2*, Genetic testing, Ovarian cancer