

Exploring the role of homologous recombination deficiency and BRCA1/2 mutations in endometrial cancer

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- 1. Endometrial cancer is part of the gBRCA1/2-associated hereditary breast and ovarian cancer (HBOC) syndrome. *This thesis*
- 2. A typical gBRCA1/2-associated endometrial carcinoma is TP53 mutated and is of high-grade. This thesis
- 3. Homologous recombination deficiency frequently occurs in *TP53* mutated endometrial carcinomas, giving rise to new treatment opportunities. *This thesis*
- 4. Tumor-first genetic testing for hereditary cancer syndromes is more efficient and should become standard clinical practice. *This thesis*
- 5. Now that molecular characterization of cancers is becoming more and more relevant for diagnostic subclassification of tumors and for precision medicine, pathologist should be trained as morpho-molecular pathologist. *Inspired by Moore et al. J Clin Pathol. 2018;71(4):285-290.*
- 6. When implementing tumor-first genetic testing as a routine part of pre-screening for hereditary variants, a multidisciplinary approach with a good collaboration between the pathology department and the clinical genetics department is essential to prevent interpretive errors of detected variants.
- 7. The opinion of the *BRCA1/2* mutation carrier should be considered when making a decision on the extent of risk-reducing surgery, and whether or not to extent the risk-reducing salpingo-oophorectomy with a hysterectomy ("shared decision-making").
- 8. An increased risk is not the same as a high risk. Researches should be encouraged also to present absolute measures (e.g. absolute risks).
- 9. "Absence of evidence does not equal evidence of absence." Adapted from *Carl Sagan*, 1997.
- 10. People should more often be uncomfortably uncertain than comfortably wrong. Adapted from *Ozan Varol "think like a rocket scientist"*, 2020.
- 11. In alle dagen zit goud verstopt. Adapted from Maris A.M, "geef me je echt", 2021