

p53 immunohistochemistry on 39.4% with a sensitivity of 98.5% to detect p53abn (99.6% negative predictive value). Cytologic features including tumor giant cells, smudged chromatin, cherry-red/macronucleoli, and atypical mitoses accurately predicted p53abn. In 7/292, p53abn upgraded ESGO risk groups (2 to intermediate-risk, 5 to high-risk). EEC12/stage IA patients had an excellent cause-specific 5-year survival of 98.5%.

**Conclusions** Pathologists can select cases for p53 testing with high sensitivity and low risk of false negativity. Molecular characterization of endometrial carcinomas has great potential to refine ESGO risk classification for a small subset but offers little value for approximately half of endometrial carcinomas, namely, EEC12/stage IA.

EPV104/#228

### EMERGING IMMUNOTHERAPY PARADIGMS IN ADVANCED ENDOMETRIAL CANCER: THE EFFECT OF ONLINE EDUCATION ON CLINICIAN KNOWLEDGE AND CONFIDENCE

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**Objectives** This study determined whether online continuing medical education (CME) could improve the knowledge of oncologists (oncs) and obstetricians/gynaecologists (obs/gyns) regarding the rationale and evidence for immunotherapy paradigms in advanced endometrial cancer.

**Methods** A 30-minute online video lecture was launched for physicians outside the USA August 2020 with data collected to November 2020. Educational effect assessed with repeated-pairs pre-/post-activity- individual participants serving as own control. 3 multiple-choice, knowledge questions and 1 self-efficacy, 5-point Likert scale confidence question were analyzed. Chi-squared test assessed pre- to post-activity change (5% significance level,  $P < .05$ ). Magnitude of change in total number of correct responses overall, and for each question, determined with Cramer's V ( $< .06$ =Modest,  $0.06$ – $0.15$ =Noticeable,  $0.16$ – $0.26$ =Considerable,  $> .26$ =Extensive).

**Results** 142 obs/gyns and 60 oncs completed pre- and post-activity questions. Positive educational effect was observed for obs/gyns (noticeable effect,  $V = .092$ ,  $P < .01$ ; average% of correct responses increasing from 33 to 42%) and oncs (noticeable effect,  $V = .150$ ,  $P = .0043$ ; average% of correct responses increasing from 47 to 62%). Increases in correct responses post-activity seen for questions on response to 2nd line chemotherapy (% relative improvement, obs/gyn: 23%, oncs: 22%), rationale for immunotherapy (obs/gyns: 24%, oncs: 72%), data for the dostarlimab GARNET trial (obs/gyns: 36%, oncs: 21%). Confidence in knowledge of the evidence for immunotherapy strategies increased post-activity (total average confidence shift: 27% obs/gyns and 40% oncs). Overall, 22% of learners' responses were improved and 39% of learners' responses were reinforced.

**Conclusions** This online CME activity resulted in a positive educational impact for both clinical specialties. However, education gaps remained evident post-activity.

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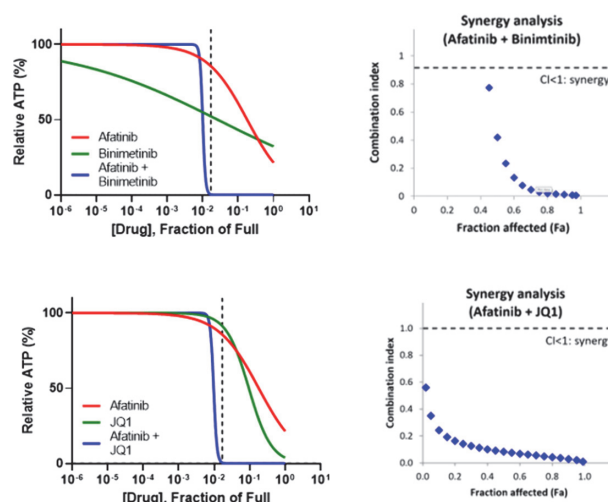
### COMBINATION TARGETED TREATMENT WITH MEK AND PAN-ERBB INHIBITORS ENHANCES ANTITUMOR ACTIVITY IN ERBB AMPLIFIED EX-VIVO SEROUS ENDOMETRIAL CANCER CELLS

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**Objectives** ERBB pathway alterations present therapeutic targets in high grade endometrial cancer (EC), but efficacy can be limited by persistent co-activation of other ERBB binding partners. The efficacy of dual-inhibition MEK+pan-ERBB or BET+pan-ERBB in an ERBB2/ERBB3 amplified EC was investigated via 3D microcancer ex-vivo cell assay.

**Methods** Tumor was prospectively collected from a patient with stage IIIc1 serous EC. Whole exome, mRNA, and Mate-Pair genomic characterization was performed. Tumor cells were grown in 3D culture and subjected to titrating drug treatments. Cell viability was determined by the CellTiter-Glo Luminescent Assay. Data transformation and dose-response curves were generated using GraphPad PRISM using the variable slope model. CalcuSyn software with the Chou-Talalay method analyzed drug interactions and synergy. Afatinib, binimetinib, and JQ1 were used to inhibit pan-ERBB, MEK1/2, BET, respectively. For translational relevance, inhibitory effect was defined as percent reduction in ATP from baseline at the



**Abstract EPV105/#237 Figure 1** Microcancer ex vivo exposure to MEK+pan-ERBB inhibitors. Dose response curves of single and combination treatments (left) were 10-fold titrated across 8 log doses for each agent. The highest concentration (i.e. fraction of ful (FoF)=1) of afatinib, binimetinib and afatinib+JQ1 was 3  $\mu$ M, 10  $\mu$ M, and 50  $\mu$ M, respectively. The physiologically achievable concentration of afatinib is insicated (dotted lines). A combination index (CI, right) was used to assess synergy with afatinib+binimetinib and afatinib+JQ1 as shown by Fa-CI plots.