

($p=0.192$). Disease progression and overall survival did not differ between groups ($p=0.537$, $p=0.465$, respectively). However, patients operated with MIS had almost 3 times risk to recur at their vaginal cuff or pelvis (Odds Ratio (OR) 95% Confidence Interval (CI) 2.80 (1.80–4.36)). In a multivariable analysis, including age, comorbidities, disease stage, CA-125 and lymph-vascular space invasion, MIS was associated with an increased risk for local (vaginal cuff or pelvic) recurrence (OR 95% CI 3.30 (1.69–6.48)).

Conclusions In women with HGEC, MIS was associated with higher rates of local recurrence as compared to laparotomy

OP013/#417 COST-EFFECTIVENESS OF DOSTARLIMAB IN ADVANCED RECURRENT DEFICIENT MISMATCH REPAIR ENDOMETRIAL CANCER PATIENTS

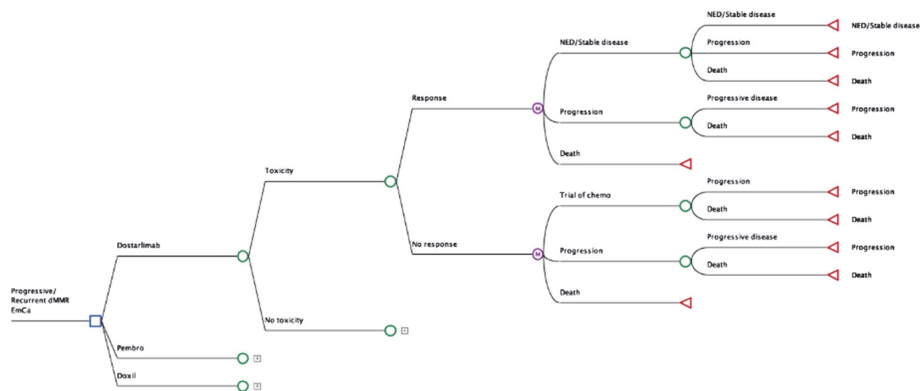
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10.1136/ijgc-2021-IGCS.30

Objectives Women with recurrent endometrial cancer who fail carboplatin and paclitaxel have a poor prognosis with few effective options. The recent GARNET Trial showed promising results for dostarlimab in these patients. We developed a decision model to compare the cost-effectiveness of dostarlimab to other treatment options in patients with progressive/recurrent deficient mismatch repair (dMMR) endometrial cancer who have failed first-line chemotherapy.

Methods A Markov model was created to simulate the clinical trajectory of women with progressive/recurrent dMMR endometrial cancer who failed carboplatin and paclitaxel (figure 1). The initial decision point in the model was treatment with either dostarlimab, pembrolizumab or pegylated liposomal doxorubicin (PLD). Model probabilities, cost and utility values were derived with assumptions drawn from published literature. The effectiveness was measured in terms of quality adjusted life years (QALYs) gained. The primary outcome was incremental cost-effectiveness ratios (ICERs), expressed in 2018 US dollars/QALYs. One-way sensitivity analyses were performed to vary the assumptions across a range of plausible values.

Results PLD was the least costly strategy at \$54,307, followed by pembrolizumab (\$160,780) and dostarlimab (\$251,132). PLD was cost-effective compared with



Abstract OP013/#417 Figure 1

Abstract OP013/#417 Table 1 Outcomes of study population

Treatment	N	Response Rate (%)	Total Costs (\$)	QALYs	ICER
PLD	2,000	10	108,615,907	1,102	N/A
Pembrolizumab	2,000	29	321,561,768	1,876	Extended dominance
Dostarlimab	2,000	42	502,265,891	3,074	199,621

Assuming a population of 6,000 progressive/recurrent dMMR endometrial cancer patients

dostarlimab with an ICER of \$199,621, while pembrolizumab was subjected to extended dominance (table 1). Multiple one-way sensitivity analyses did not substantially impact the cost-effectiveness.

Conclusions Dostarlimab is associated with greater survival compared with other treatments for women with recurrent dMMR endometrial cancer. However, the agent is substantially more costly.

OP014/#480 TARGETED MOLECULAR TESTING IN ENDOMETRIAL CARCINOMA: VALIDATION OF A RESTRICTED TESTING PROTOCOL

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10.1136/ijgc-2021-IGCS.31

Objectives The World Health Organization (WHO) endorsed molecular classification of endometrial carcinoma (EC) to be incorporated in routine diagnostic workup, by evaluating p53 and mismatch repair (MMR) protein immunohistochemistry (IHC), as well as pathogenic mutations in the gene encoding DNA polymerase epsilon (POLE). The latter is currently the least affordable or accessible step. We investigated whether POLE testing can be omitted in patients who based on stage, grade and lymphovascular space invasion (LVI) criteria would not usually be directed to adjuvant therapy.

Methods Using data from a single cancer centre (n=460) in Vancouver, and a population-based cohort in Tubingen (n=452), we compared the WHO recommended molecular testing of the entire cohort (n=912) with a restricted protocol: p53 and MMR IHC on all cases, but POLE sequencing

not done on 'very low-risk' ECs, defined as Stage 1A, G1/G2, no LVI, MMR proficient and without p53 abnormalities.

Results 30% of full cohort and 38% of population-based patients were classified as 'very low-risk', and did not undergo POLE testing. 'Very low-risk' ECs with unknown POLE status showed excellent clinical outcomes in both univariable and multivariable survival models. Amongst G1/G2 EEC, 14/566 (2.5%) were p53abn, and G1/G2 EEC constituted 14/166 (8.4%) of all p53abn ECs.

Conclusions Molecular classification of EC can be safely and more pragmatically incorporated into routine clinical practice using universal MMR and p53 IHC, and foregoing POLE testing in 'very low-risk' ECs where this has no therapeutic impact. Restricting molecular testing to high-grade/high-risk EC would miss some p53abn patients.

OP015/#492

FURTHER STRATIFICATION OF NO SPECIFIC MOLECULAR PROFILE (NSMP/P53WT) ENDOMETRIAL CARCINOMAS TO REFINE PROGNOSIS AND IDENTIFY THERAPEUTIC OPPORTUNITIES

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10.1136/ijgc-2021-IGCS.32

Objectives Molecular classification identifies >50% of endometrial cancers (ECs) as having 'no specific molecular profile/NSMP'; without mismatch repair deficiency, p53 IHC abnormalities, or pathogenic POLE mutations. Clinical presentation and outcomes within NSMP ECs are diverse and optimal treatment unclear with new ESMO/ESTRO/ESP guidelines unchanged for this molecular subtype. Better biomarkers are needed to predict if and what adjuvant therapies are needed.

Methods We characterized the clinicopathological and molecular (IHC+NGS) profiles of 1047 NSMP ECs in women from population-based and institutional cohorts, testing for associations with treatment response and outcomes.

Results Key pathologic and molecular features associated with survival parameters ($p < 0.01$) are tabulated below. 31% of NSMP ECs had CTNNB1 mutations, however, associations with outcomes (PFS) were observed only within Gr1/2 early-stage endometrioid ECs ($p = 0.03$), or if restricted to ECs without substantial LVI or L1CAM overexpression ($p < 0.005$). TP53 mutations (with normal p53IHC) were discovered in 41 women with a trend ($p = 0.06$) to worse survival. On

Abstract OP015/#492 Table 1

	Full cohort	Stage IA	Stage IB	St II-IV
Total	1047 (100%)	673 (67%)	164 (16%)	175 (17%)
Age -median (range)	61 (22-96)	59 (22-94)	66(32-93)	63(32-96)
BMI-median (range)	31 (16-82)	31 (16-82)	31 (16-66)	29 (16-53)
Grade 1/2/3 (%)	69/17/14%	79/15/7%	66/19/15%	34/29/37%
Histo endometrioid	974 (93%)	641 (96%)	159 (97%)	142 (82%)
LVI	82/15/3%	93/6/1%	73/20/6%	45/45/10
neg/focal/substantial				
ESMO low/int/HR/high	52/10/14/2 3	78/0/17/5	0/68/17/15	0/0/0/94
ER neg/weak/strong	7/16/77	5/14/81	7/19/73	17/24/59
PR neg/weak/strong	11/20/69	8/18/74	5/29/66	31/19/50
L1CAM overexpression	15%	13%	13%	24%
ARID1A loss (IHC)	33%	31%	38%	36%
PIK3CA mutation	39%	39%	32%	35%
Post-surgical Rx (%)	65/14/9/12	83/11/4/2	38/39/14/9	21/5/25/47
none/VB/EBRT/chemo+/- RT				
Overall survival event	198 (19.5%)	79(12%)	38(24%)	64 (37%)
Disease specific survival event	82 (8%)	17 (3%)	11 (7%)	43 (26%)
Progression/recurrence events	102 (10.4%)	37 (5.7%)	19 (12%)	44 (29%)

Cox regression and Kaplan-meier analysis demonstrate statistically significant associations w/outcomes (OS, DSS, PFS) ($p < 0.01$) for all parameters except Age, BMI, and PIK3CA (OS +DSS only) + ARID1A (OS only)

multivariable analysis only grade (3vs.1/2) maintained significance. 8% of this cohort would be eligible for current molecular classification de-escalation trials. Treatment received did not impact survival within low-, intermediate-, or high-intermediate risk NSMP ECs. Within high-risk, the most favorable outcomes were observed in women who received pelvic radiation with no observed benefit of chemotherapy.

Conclusions Additional prognostic stratification of NSMP ECs can be achieved with both pathologic and molecular features. Further study within NSMP subgroups may identify conventional, hormonal or targeted therapies that are more effective.

OP016/#563

MORBIDITY AND QUALITY OF LIFE OF SENTINEL LYMPH NODE MAPPING IN ENDOMETRIAL CANCER. INTERIM ANALYSIS OF A PROSPECTIVE RANDOMIZED TRIAL (ALICE TRIAL)

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10.1136/ijgc-2021-IGCS.33

Objectives Despite the growing evidence of sentinel lymph node mapping (SLN) in endometrial cancer, studies addressing morbidity and impact on quality of life (QoL) are still scarce. Our aim was to evaluate treatment morbidity and QoL for SLN ± systematic lymph node dissection (LND) in endometrial cancer.