

Methods: Patients with platinum-sensitive (cohort 1) and platinum-resistant (cohort 2) aOC were enrolled. Patients with known/suspected deleterious germline *BRCA1/2* mutation and ≥ 2 prior lines of chemotherapy were eligible. The primary endpoint was objective response rate (ORR) assessed by an independent review committee (ORR_{IRC}) per RECIST v1.1.

Results: As of 2 Feb 2020, 113 pts (cohort 1, n=90; cohort 2, n=23) were enrolled. Median age was 54 yr (range: 34-79), 25.6% (n=29) of pts had received ≥ 4 prior systemic chemotherapy lines, and 54.0% (n=61) of pts had an ECOG score of 1 at study entry. At data cutoff, median follow-up was 12.2 mo (range: 0.2-21.5). Across both cohorts, pamparib showed preliminary antitumor activity (Table). In cohort 1, confirmed ORR_{IRC} was 64.6%, median DoR was 14.5 mo (95% CI, 11.1-NE), progression-free survival (PFS) was 15.2 mo (95% CI, 10.35-NE), and median overall survival (OS) was not yet mature. In cohort 2, confirmed ORR_{IRC} was 31.6%, median DoR was 11.1 mo (95% CI, 4.21-NE), median PFS was 6.2 mo (95% CI, 4.11-NE), and median OS was 13.6 mo (95% CI, 7.13-NE). Overall, the most common treatment-related AE was anemia (any grade, 89%; grade ≥ 3 , 42%); following a per-protocol proposed dose modification algorithm, the incidence of grade ≥ 3 anemia was reduced to 25.6%.

Table: 820P		
Efficacy-evaluable population	Cohort 1 (n=82)	Cohort 2 (n=19)
Best overall response, n (%)		
Complete response	8 (9.8)	0 (0)
Partial response	45 (54.9)	6 (31.6)
Stable disease	25 (30.5)	12 (63.2)
Progressive disease	4 (4.9)	1 (5.3)
Objective response rate, % (95% CI)		
Confirmed	64.6 (53.3-74.9)	31.6 (12.6-56.6)
Disease control rate, % (95% CI)	95.1 (88.0-98.7)	94.7 (74.0-99.9)
Clinical benefit rate, % (95% CI)	74.4 (63.6-83.4)	52.6 (28.9-75.6)
Time to response, median mo (range)	1.68 (1.3-6.3)	1.38 (1.2-1.4)
Duration of response, median mo (95% CI)	14.5 (11.1-NE)	11.1 (4.21-NE)
PFS, median mo (95% CI)	15.2 (10.35-NE)	6.2 (4.11-NE)

Abbreviations: NE, not estimable; PFS, progression-free survival.

Conclusions: Promising antitumor activity was observed in pts with platinum-sensitive/resistant aOC. Pamparib was generally tolerated, with no new safety signals. Pamparib is being evaluated as monotherapy and combination therapy for other solid tumors.

Clinical trial identification: NCT03333915.

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821P Timing of adverse events during maintenance treatment with rucaparib for recurrent ovarian cancer in the phase III ARIEL3 study

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Background: Maintenance treatment with the poly (ADP-ribose) polymerase (PARP) inhibitor rucaparib significantly improved progression-free survival vs placebo regardless of biomarker status in patients with recurrent ovarian cancer in the phase III ARIEL3 study (NCT01968213). This exploratory analysis evaluated the timing of treatment-emergent adverse events (TEAEs).

Methods: Patients were randomised 2:1 to receive oral rucaparib (600 mg twice daily) or placebo until disease progression, unacceptable toxicity or other reason for discontinuation. Time to onset of TEAEs was assessed.

Results: Of the 564 patients randomised in ARIEL3, 561 patients (372 for rucaparib and 189 for placebo) were included in the safety population (updated visit cut-off date, 31 December 2017). Median duration of treatment was 8.3 months for rucaparib and 5.5 months for placebo. Overall, 15.3% of patients receiving rucaparib and 1.6% of patients receiving placebo discontinued treatment due to TEAEs (excluding disease progression). Safety data, including median time to onset of the first event, for TEAEs occurring in $\geq 35\%$ of patients are presented in the table.

Conclusions: In the rucaparib group, median time to first onset for most of the frequently reported non-haematological TEAEs was within 1 month of therapy. In the rucaparib and placebo groups, median time to first onset of anaemia/decreased haemoglobin occurred within 3 and 2 months of therapy, respectively.

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Table: 821P						
TEAE	Patients with TEAE, n (%)		Patients with dose reduction or treatment interruption due to TEAE, n (%)		Median time to first onset, days (95% CI)	
	Rucaparib	Placebo	Rucaparib	Placebo	Rucaparib	Placebo
Nausea	282 (75.8)	69 (36.5)	56 (15.1)	2 (1.1)	5 (4-6)	17 (7-57)
Asthenia/fatigue ^a	263 (70.7)	84 (44.4)	46 (12.4)	6 (3.2)	15 (15-20)	30 (16-37)
Dysgeusia	148 (39.8)	13 (6.9)	5 (1.3)	0	10 (6-15)	11 (3-29)
Anaemia/decreased haemoglobin ^a	145 (39.0)	10 (5.3)	66 (17.7)	1 (0.5)	67 (57-85)	54 (15-140)
Constipation	141 (37.9)	46 (24.3)	10 (2.7)	1 (0.5)	40 (29-55)	46 (23-81)
Vomiting	138 (37.1)	29 (15.3)	35 (9.4)	2 (1.1)	24 (15-39)	84 (36-129)

^aCombined terms.

TEAE, treatment-emergent adverse events.

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822P Comparison of toxicities of PARP inhibitors used in gynaecological cancers observed at a large cancer centre

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Background: Olaparib (O) was first poly ADP-ribose polymerase inhibitor (PARPi) approved for use in the UK as maintenance therapy in relapsed BRCA mutated ovarian cancer in 2017. Since then Niraparib (N) and Rucaparib (R) have also been approved as maintenance therapy in patients with relapsed ovarian cancer with and without BRCA gene mutations. Though clinicians now have three drugs to choose from, a lack of head-to-head data means the choice is not a straight forward one. We aim to compare toxicities of the PARPi drugs in order to give clinicians a better idea of which drugs to use in certain scenarios. We report the real world experience of the three PARPi at a UK cancer centre.

Methods: Retrospective data was collected from every patient with a gynaecological cancer who had a PARPi up to the 19th of March 2020. Data lock was the 10th of May 2020.

Results: 119 patients with relapsed platinum-sensitive ovarian/fallopian tube or primary peritoneal cancer were included. The group contained 72 patients with BRCA wild-type (wtBRCA), 46 patients with germline mutated BRCA (gBRCA) and one patient with an unknown BRCA status. 71 patients received N, 37 received O and 12 patients received R. gBRCA patients were more likely to receive O. wtBRCA patients were more likely to receive N or R. The median number of cycles was 5. The preliminary median PFS for the gBRCA population is 14.9 months. The preliminary median PFS for the wtBRCA population is 7.2 months. Treatment interruption, dose reduction and early stoppage all favoured O. The incidence of grade ≥ 3 cytopenia was similar across the three drugs. Derangement of liver function, abdominal pain, nausea, vomiting and diarrhoea was observed more often with R. Hypertension, insomnia, mucositis and dyspnoea were seen with more often with N. Evaluation of correlation between BRCA status and incidence of adverse events is ongoing. 41 patients have thus far completed chemotherapy post progression on PARPi. The myelosuppressive effects of chemotherapy post-PARPi and its correlations with the individual drugs and BRCA status is being evaluated. Mature data will be presented at the meeting.

Conclusions: Efficacy of PARPi was superior in the gBRCA population compared to wtBRCA. O was the least toxic PARPi.

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823P Extended follow-up of a real-world cohort of patients (pts) with BRCA mutation (BRCAm) relapsed epithelial ovarian cancer (EOC) receiving olaparib maintenance therapy: The GINECO RETROLA study

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Background: Olaparib was initially approved by the European Medicines Agency (EMA) as maintenance treatment for pts with BRCAm platinum-sensitive relapsed high-grade EOC, following the results of a randomized phase II trial (study 19). The RETROLA study aimed to evaluate whether the outcome observed in this clinical trial are reflected in routine clinical practice, in a real-world cohort of pts.

Methods: We planned to include 130 pts in this retrospective cohort. French centers (n=28) representative of French regions and of mode of practice were asked to