

## Esmo preview - Novel checkpoints add to the Parp showdown



[Jacob Plieth](#)

While the texts of oral and late-breaking Esmo abstracts are under wraps until next week, today's unveiling of the meeting's poster presentations has given investors a few more showdowns to look forward to. These include AstraZeneca and Pfizer's Ox40 agonists, and the leading checkpoint inhibitors in urothelial carcinoma.

The data add to Esmo's most keenly awaited confrontations, including those between Clovis, Medivation, Astra and Tesaro's Parp inhibitors (see table below). The top places at the meeting's late-breakers have also been confirmed, though bizarrely one of the most important studies has been edged out of the press programme by Oncogenex's failed Affinity trial.

That prostate cancer study, the latest in a litany of flops for Oncogenex's custirsen, [failed in August](#) despite an earlier protocol change. Yet it features not only as a late-breaker at the third of Esmo's presidential sessions, but also in the special press programme on Monday.

Hopefully Esmo, which in previous years has given special treatment to failures like GlaxoSmithKline's Mage-A3 vaccine, is encouraging lessons to be learned from disaster. However, Affinity's inclusion comes at the expense of Bristol-Myers Squibb's failed Checkmate-026 first-line NSCLC trial – one of the meeting's hottest tickets.

Bristol is expected to detail essential findings regarding Opdivo's efficacy in patients with different levels of PD-L1 expression ([Bristol swings for the fences and strikes out, August 5, 2016](#)). Two of Opdivo's checkpoint inhibitor competitors, Merck & Co's Keytruda and Roche's Tecentriq, star in their own NSCLC late-breakers, though the latter's Oak trial is in second-line use.

### Hot Parps

Failure also features in the area of Parp inhibition, where the [Gold trial of AstraZeneca's Lynparza in advanced gastric cancer](#) has a Saturday late-breaker spot.

After Pfizer's \$14bn takeout of Medivation interest in Parp inhibitors has grown, and investors will also be focusing on data to be presented at Esmo from the [Ariel2 study of Clovis's rucaparib](#), and the [Nova trial of Tesaro's niraparib](#), in ovarian cancer.

Other mechanisms of interest include CDK4/6 inhibition, where Novartis's ribociclib is squaring up to Pfizer's marketed breast cancer drug Ibrance. The Esmo late-breaker concerns first-line use of ribociclib plus letrozole in HR-positive, Her2-negative breast cancer – likely the [Monaleesa-2 study that was halted early for efficacy in May](#).

### Selected Esmo 2016 presentations

Date	Project	Company	Study	Trial ID	Abstract
7 Oct	KTE-C19	Kite	Zuma-1	NCT02348216	1048O
7 Oct	Selinexor	Karyopharm	-	NCT02025985	854O
7 Oct	Prexasertib	Lilly/Array	-	NCT02203513	855O
7 Oct	Rucaparib	Clovis	Ariel2	NCT01891344	856O
7 Oct	ANG1005	Angiochem	-	NCT02048059	324O
8 Oct	Ribociclib	Novartis	Monaleesa-2	NCT01958021	LBA1_PR
8 Oct	Opdivo	Bristol-Myers Squibb	Checkmate-275	NCT02387996	LBA31_PR
8 Oct	Keytruda	Merck & Co	Keynote-052	NCT02335424	LBA32_PR
8 Oct	Niraparib	Tesaro	Nova	NCT01847274	LBA3_PR
8 Oct	Lynparza	AstraZeneca	Gold	NCT01924533	LBA25
8 Oct	Talazoparib	Pfizer (Medivation)	-	NCT02282345	153PD
8 Oct	CRLX101	Cerulean	-	NCT02389985	864P
9 Oct	Opdivo	Bristol-Myers Squibb	Checkmate-026	NCT02041533	LBA7_PR
9 Oct	Keytruda	Merck & Co	Keynote-24	NCT02142738	LBA8_PR
9 Oct	Opdivo	Bristol-Myers Squibb	Checkmate-141	NCT02105636	LBA4
9 Oct	Tecentriq	Roche	Oak	NCT02008227	LBA44_PR
9 Oct	Xtandi	Medivation/Pfizer	Premiere	NCT02288936	726PD
9 Oct	NY-ESO-1	Adaptimmune	-	NCT01343043	1075P
9 Oct	Ipafricept	Bayer/Oncomed	-	NCT02050178	367PD
9 Oct	Cometriq + Opdivo	Exelixis/BMS	-	NCT02496208	774PD
9 Oct	Lirilumab + Yervoy	Innate Pharma/BMS	-	NCT01592370	1086P
10 Oct	Custirsen	Oncogenex	Affinity	NCT01578655	LBA9_PR
10 Oct	Cotellic + Tecentriq	Exelixis/Roche	-	NCT01656642	1109PD
10 Oct	Epacadostat + Keytruda	Incyte/Merck & Co	Keynote-037	NCT02178722	1110PD
10 Oct	Avelumab	Pfizer/Merck KGaA	Javelin Solid Tumor	NCT01772004	777PD
10 Oct	MEDI0562	AstraZeneca	-	NCT02318394	1052PD
10 Oct	PF-04518600	Pfizer	-	NCT02315066	1053PD

Among the abstracts that have already been made available, Pfizer and Merck KGaA's avelumab is contesting the first approved use of Roche's Tecentriq - urothelial carcinoma. Early data relate only to 109 patients with over four months' follow-up, showing three complete and 15 partial remissions.

Opdivo and Keytruda are making challenges of their own in this cancer type - their respective Checkmate-275 and Keynote-052 trials have late-breaker slots. And data will also be presented from Opdivo's Checkmate-141 head and neck cancer trial, [halted for efficacy in January](#) before swift US and EU filings in this cancer.

Of course, Esmo is not all about PD-1/PD-L1 inhibitors. Among the unveiled abstracts is a first-in-human trial of Pfizer's Ox40 agonist, PF-04518600, showing just one partial remission (PR) in 25 patients, while Astra's rival MEDI0562 has shown one PR in 19 evaluable patients.

Elsewhere, Incyte's IDO1 inhibitor epacadostat has shown a 58% ORR in 19 first-line melanoma patients in the

Keynote-037 combination trial with Keytruda. Other trials suggest the viability of combining Innate Pharma's lirilumab with Bristol's Yervoy or Opdivo, though the Esmo abstract mainly details the combos' toxicity profiles.

Still, the much bigger event for Innate investors is the monotherapy trial Effikir, in AML, whose results are imminent. Likewise, Kite Pharma investors now know that full data from the Zuma-1 interim readout toplined this week will not come until the ASH meeting in December, so the Esmo update of Zuma-1's phase I portion might be purely academic.

*Robin Davison will report from the Esmo meeting on October 7-11. He can be followed on [@RobinDavison2](#). EP Vantage has also produced a free [backgrounder](#) ahead of the meeting, highlighting recent breakthroughs, setbacks and approaching data points.*

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