

Supplementary Material

A Phase Ib Study Assessing the Safety, Tolerability, and Efficacy of the First-in-Class Wee1 Inhibitor Adavosertib (AZD1775) as Monotherapy in Patients with Advanced Solid Tumors

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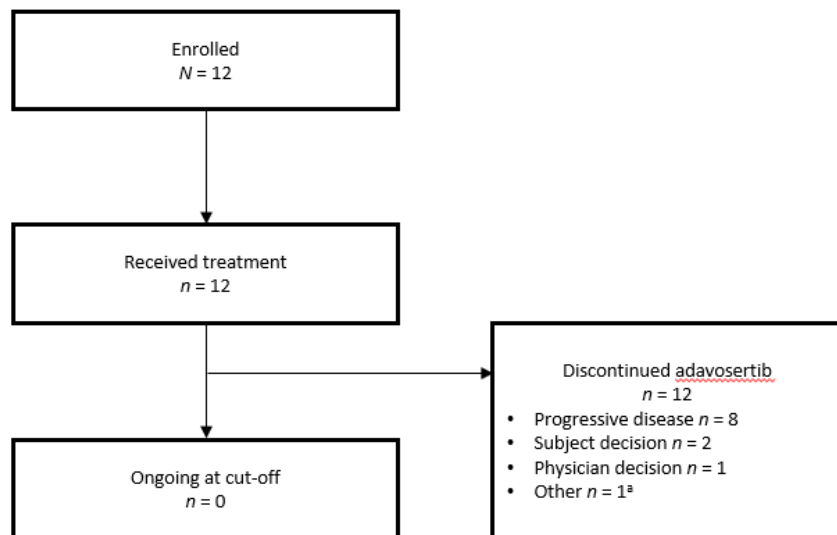
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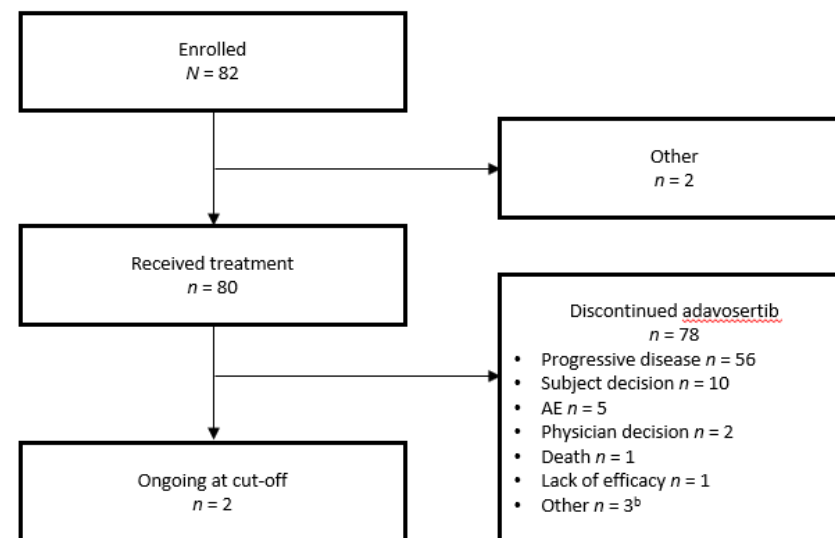
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Supplementary Figure S1. Patient disposition across Part A (safety lead-in) and Part B (expansion cohorts)

Part A

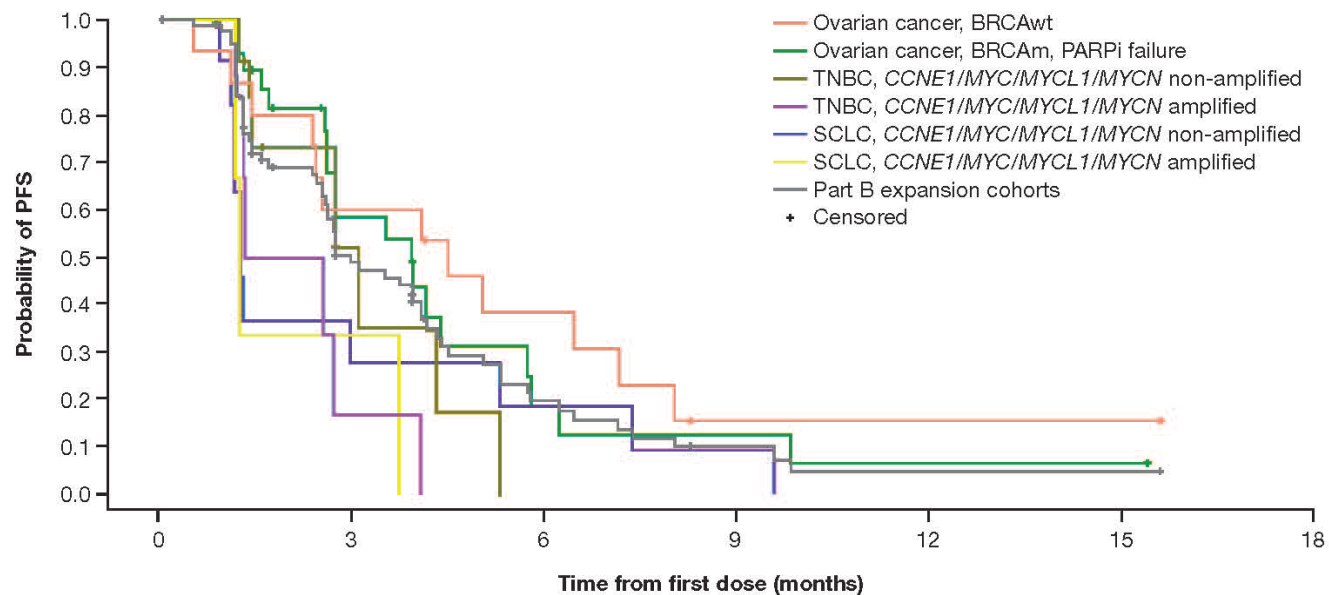


Part B



^aLost to follow-up as a result of issues with insurance; ^bOne patient in the TNBC, biomarker-negative cohort (increased worsening of AEs and increase in tumoral markers) and two in the ovarian cancer BRCAm, PARPi failure cohort (clinical progression). AE adverse event, BRCAm breast cancer gene 1/2 mutation, N number of patients enrolled, n number of patients, PARPi poly(ADP-ribose) polymerase inhibitor, TNBC triple-negative breast cancer

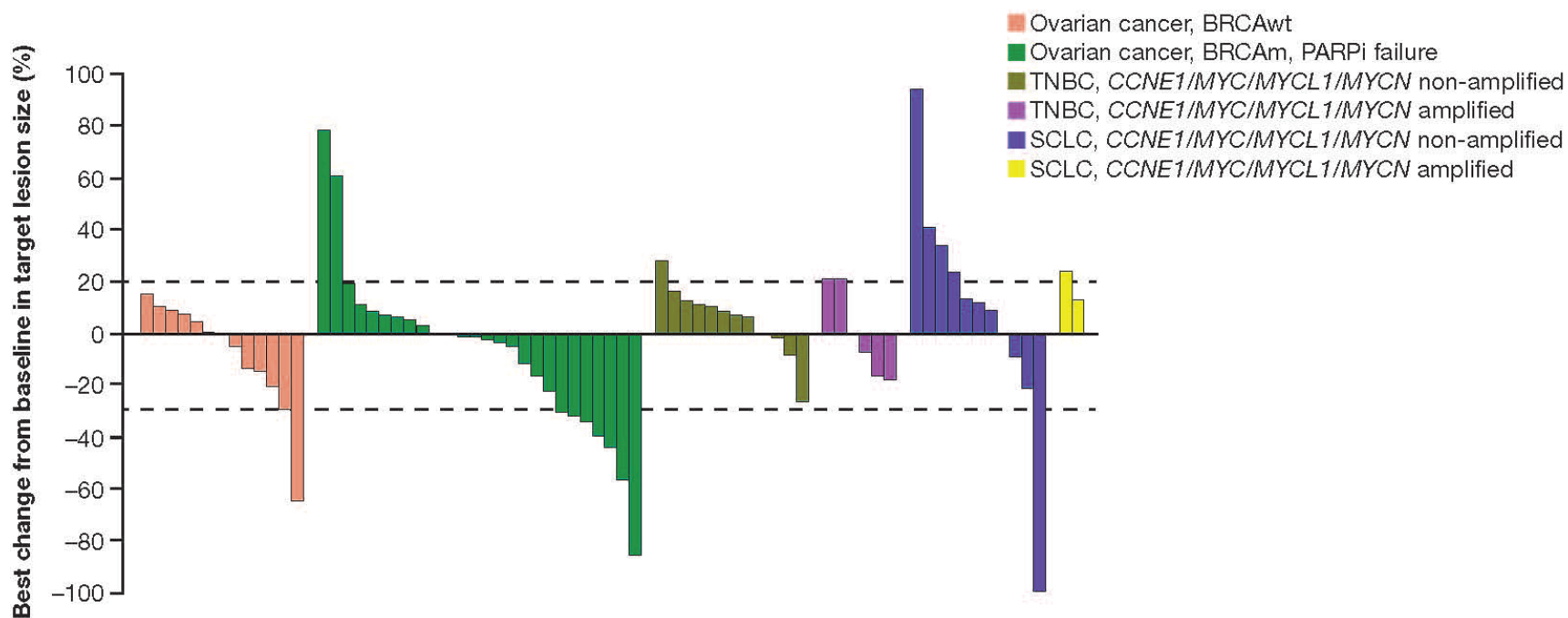
Supplementary Figure S2. Progression-free survival (PFS) across all tumor types



	0	3	6	9	12	15	18
Ovarian cancer, BRCAwt: 16/12*	16	9	5	1	1	1	0
Ovarian cancer, BRCAm, PARPi failure: 30/19*	30	12	3	2	1	1	0
TNBC, CCNE1/MYC/MYCL1/MYCN non-amplified: 13/8*	13	3	0				
TNBC, CCNE1/MYC/MYCL1/MYCN amplified: 6/6*	6	1	0				
SCLC, CCNE1/MYC/MYCL1/MYCN non-amplified: 12/11*	12	3	2	1	0		
SCLC, CCNE1/MYC/MYCL1/MYCN amplified: 3/3*	3	1	0				
Part B expansion cohorts: 80/59*	80	29	10	4	2	2	0

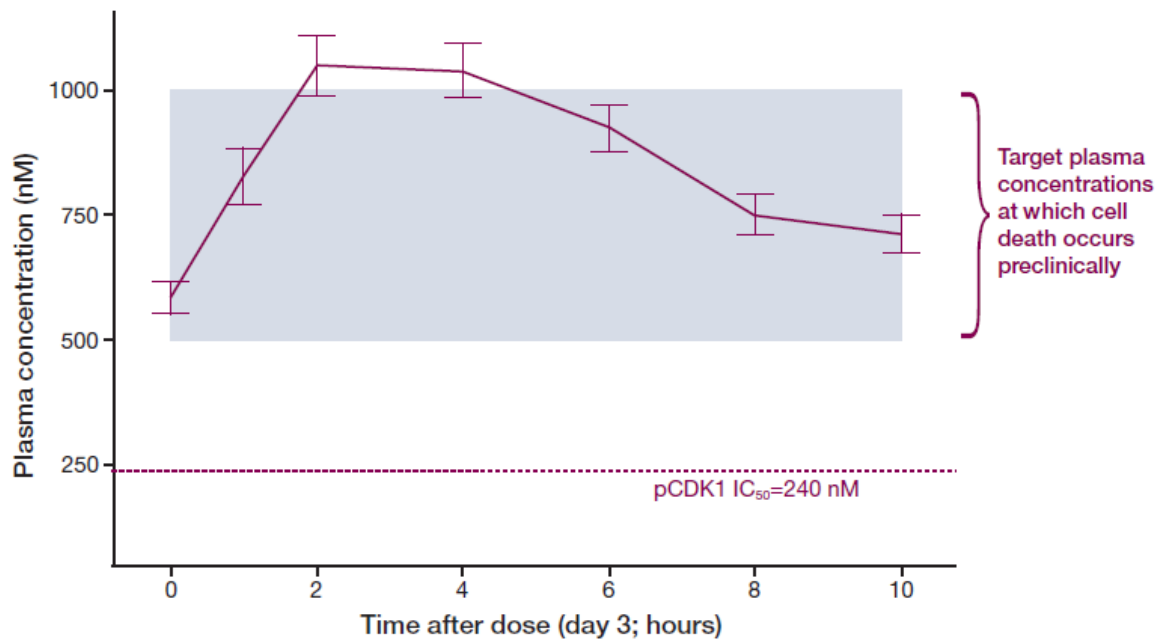
*Number of patients in the full analysis set/number of events. *BRCAm* breast cancer gene 1/2 mutation, *BRCAwt* breast cancer gene 1/2 wild type, *CCNE1* cyclin E1, *PARPi* poly(ADP-ribose) polymerase inhibitor, *SCLC* small-cell lung cancer, *TNBC* triple-negative breast cancer

Supplementary Figure S3. Waterfall plot of target lesion size and best percentage change from baseline in all cohorts (full analysis set)



RECIST version 1.1 partial response: $\geq 30\%$ decrease in the sum of diameters of target lesions, taking as reference the baseline sum diameters; progressive disease: $\geq 20\%$ increase in the sum of diameters of target lesions, taking as reference the smallest sum on study. *BRCAm* breast cancer gene 1/2 mutation, *BRCAwt* breast cancer gene 1/2 wild type, *CCNE1* cyclin E1, *PARPi* poly(ADP-ribose) polymerase inhibitor, *SCLC* small-cell lung cancer, *TNBC* triple-negative breast cancer

Supplementary Figure S4. Clinical pharmacokinetics



Plasma concentrations of adavosertib at 175 mg bid (day 3 exposure, geometric mean \pm standard deviation). *bid* twice a day, IC_{50} half-maximal inhibitory concentration, *pCDK1* phosphorylated cyclin-dependent kinase 1