		Grade 1	Grade 2	Grade 3	Grade 4	Grade 5
Endocrine				-		-
	Adrenal insufficiency			2		
	Hyperthyroidism / decreased TSH	7	7	1		
	TSH	1	13			
Gastrointe	stinal					
	Abdominal pain	1	1	1		
	Colitis		1	3		
	Diarrhea	9	2	4		
	Hepatitis			1		
	Nausea	7	3			
	Pancreatitis			1		
	Vomiting	3		1		
General						
	Chills	11				
	Injection site reaction	19				
	Infusion related reaction		3			
	Multi-organ failure					1
	Weight gain	1	1			
Lab investi	igations					
	Increased AST	3	1	2		
	Increased amylase	1	1	1		
Metabolisn	n and Nutrition					
	Anorexia	5				
	Hyperglycemia	1	1		1	
Musculosk	eletal and Connective Tissue					
	Arthralgias / arthritis	6	1			
	Back pain	1	2	1		
	Bone pain	2		1		
	Muscle weakness	1	1	1		
	Myalgia	5	2			
Nervous S	ystem					
	Dizziness	4	1			
	Headache	6				
	Syncope			1		
Respirator	y, Thoracic					
	Cough	6				
	Hypoxia			1		
Skin and S	Subcutaneous					
	Pruritus	3		1		
	Rash maculo-papular	4	1	1		
Vascular						
	Thromboembolic event (pulmonary embolism)					

**Supplemental Table 1:** <u>Adverse Events for All Study Patients</u>. All adverse events with a frequency >5%, and any adverse events with grade > grade 2, that were believed to be at least possibly related to treatment, are shown for all trial arms. The numbers represent the number of patients experiencing a particular event at any point during the treatment period, with the highest grade reported for any single individual.

Subject ID	MSI <sup>hi</sup> or HRR mutation	Best PSA change from baseline (%)	rPFS (months)	Time on trial (months)
30005 (Arm 3)	MSI <sup>hi</sup> and MSH3	-90.1	11.0	11.3
40011 (Arm 4)	MSI <sup>hi</sup>	38.2	5.3	8.2
20002 (Arm 2)	ATM	(not evaluable)	5.5	5.5
20012 (Arm 2)	ATM	22.5	0	2.5
30007 (Arm 3)	BRCA1	5.5	5.4	5.9
30008 (Arm 3)	FANCA, MLH3	32.4	2.5	3.4
30014 (Arm 3)	BRCA1	-38.0	2.6	3.0
30017 (Arm 3)	MSH2, MLH1	52.5	2.6	3.7
30018 (Arm 3)	MSH6	65.2	2.8	4.0
40009 (Arm 4)	BRCA2	24.2	1.4	1.4
40014 (Arm 4)	BRCA1	41.7	2.5	2.8

**Supplemental Table 2:** <u>HRR mutations and patient outcomes</u>. Pre-treatment plasma was available for 61/66 patients and evaluated for HRR mutations or MSI<sup>hi</sup> tumor status by cfDNA testing (Tempus). Shown are the specific mutations and outcomes (best % PSA change from baseline, rPFS, and time on trial) for patients with identified mutations.



**Supplemental Figure 1:** <u>Immunological response - IFNγ and Granzyme B fluorescent ELISPOT</u>. Peripheral blood mononuclear cells were collected from subjects at baseline, 6 weeks, 12 weeks, and 24 weeks and evaluated for antigen-specific secretion of both IFNγ and granzyme B by fluorescent ELISPOT. Shown are the spot-forming units (SFU) following stimulation with a PAP peptide library, PSA peptide library (non-specific control), or tetanus (positive control) for each patient. Patients treated in Arm 3 are colored red, and patients treated in Arm 4 are colored blue.



**Supplemental Figure 2:** Immunological response - IFNy and Granzyme B fluorescent ELISPOT. Cryopreserved peripheral blood mononuclear cells collected from 9 subjects after 24 weeks of treatment were separated into CD4, CD8 and CD14 cells by magnetic selection. CD4 or CD8 T cells were cultured with CD14 cells at a 10:1 ratio in the presence of PAP or PSA peptide libraries for 48 hours, and evaluated for granzyme B (panel A) or IFNy (panel B) secretion by ELISPOT. Shown are the spot-forming units (SFU) for each condition, with background (media only, no peptide library) subtracted from each.



**Supplemental Figure 3:** <u>Association of changes in cytokines with time on trial</u>. Time on trial was assessed with respect to changes in CXCL-10, CXCL-9, G-CSF, IFNα, IL-12p40, IL-1ß, IL-2, IL-2R, and MCP-1. For each panel, time on trial is plotted with respect to changes greater (red) or less than (blue) the median for each patient. No statistically significant differences were observed (log-rank test).



**Supplemental Figure 4**: <u>Radiographic progression-free survival correlative analyses</u>. Radiographic progression-free survival was assessed with respect to the development of grade 2 or higher immune-related adverse events (panel A), T-cell immune response to PAP (panel B), increases in serum IFNγ (panel C), and the presence of baseline HRR mutations (panel D). Statistical comparisons are made using a log-rank test, with p<0.05 considered statistically significant.