Supplementary Materials

Table S1. List of published studies with DC vaccines in gynecological and breast cancers, from 2000 to date.

Reference /	Study	Vaccine type and	DC Matur.	N°		Results: survival	
Study type	indication	preparation	stimulus	of pts	Results: immunogenicity	(PFS/OS)	AEs
Brossart-	Metastatic	* DC peptide vaccine:	TNF- α	10	* In 5 of 10 vaccinated pts,	No data	* None,
2000 [21]/	BrCa or	Her2/neu, MUC1.		(7	peptide-specific CD8+ cytolytic		particularly no
Phase I	OvCa that	* Peptide-pulsed DCs		BrCa,	T-cells were detected in the		clinically
	expressed	generated from PBMCs		3	peripheral blood after 3 Vx		relevant
	HLA-A2 and	were injected s.c. into		OvCa)	* Epitope spreading with a		anemia.
	HER-2/neu or	the upper limb close to			single tumor antigen was		* No
	MUC1.	the inguinal lymph			observed in vivo on vaccination		autoimmune
		nodes on days 1, 14, and					phenomena
		28, respectively.					observed
		* PBMCs isolated from					
		blood, were cultured in					
		medium supplemented					
		with IL-4, GM-CSF, and					
		TNF- α . DCs were					
		separately pulsed for 2					
		hours with each					
		peptide, and washed					
		before application.					
		* Peptides derived from					
		HER-2/neu (E75:					
		KIFGSLAFL, GP2:					

IISAVVGIL) and MUC1 (M1.1 STPPVHNV, M1.2: LLLLTVLTV) were synthesized * Pts received either HER-2/neu or MUC1 peptides

Triozzi-	Pts with	* DC vaccine	no maturation	10	* Biopsies of regressing lesions	* 4 days after	* Injections
2000 [20]/	metastatic	* DCs generated from		(3	showed lymphocyte infiltration	injection, regression	were well
Phase I	dermal or	monocytes obtained by		BrCa,	(TIL) associated with DCs and	of the injected tumors	tolerated.
	subcutaneous	phlebotomy and		7	necrosis	was observed in 4	* Only 1 pt
	tumors	cultured with GM-CSF		MEL)	* Injected DCs produced IFN-a	MEL pts and in 2	reported pain
	(BrCa, MEL)	and IL-4 in autologous			and expressed Fas ligand	BrCa pts	(<48h)
		plasma for 8 days			mRNA		
		* Tumors were injected			* No cytolytic activity in vitro		
		at multiple sites with 30			(reduced expression of the		
		million autologous DCs			costimulatory molecule, B7-2		
		per tumor			(CD86) on DCs after IT		
-					injection)		

Hernando-	Progressive	* DC aWTL and KLH	TNF-α	8	* A significant tumor antigen-	* A possible	* Treatment
2002 [31]/	or recurrent	vaccine.		(6	specific lymphoproliferative	correlation between	was safe,
Phase I	OvCa or	* DCs pulsed with		OvCa,	response was detected in 2 pts	the immune response	feasible and
	uterine	autologous tumor lysate		2	after two Vx	and disease	well tolerated
	sarcoma.	and KLH.		UtSar)		stabilization was	* No major
		* Crude lysates from				suggested	toxic (grade 2)
		tumor specimens (WTL)					or SAE related
		were frozen.					to the vaccine
		* PBMCs isolated from					* Minor
		leukapheresis were					general effects
		cultured for 6-7 days					included mild
		with GM-CSF.					transient
		* Inmature DCs were					fatigue, chills
		pulsed with KLH and					and low-grade
		autologous WTL in the					fever (2 pts);
		presence of GM-CSF,					no treatment
		IL-4 and TNF- α					required
		* Cells harvested in day					* No rash or
		7 or 10 were					lymphadenop
		administered i.c. in					athy or
		close proximity to the					autoimmunity
		axillary lymph nodes.					
		* pts received 3 to 23					
		injections of KLH- and					
		WTL-pulsed DC Q10D					
		or Q4W					

Vonderhei	HLA-A2-	* DC peptide hTERT	no maturation	7 (5	* hTERT-specific T lymphocytes	* PR in 1 pt was	* No
de-2004	positive pts	vaccine: autologous		ProstC	induced in 4 of 7 pts	associated with the	significant
[22]/ Phase	with	DCs loaded with hTERT		a, 2	* hTERT-specific CD8+ cells	induction of CD8+	toxicity
I	hormone-	peptide and KLH		BrCa)	after vaccination were	TILs	observed
	independent	* Generated from			identified by peptide/MHC		
	prostate	PBMCs			tetramers, proliferated, and		
	cancer or	* Matured with GM-			secreted IFN-γ after in vitro		
	refractory	CSF and IL-4, and			peptide sensitization, killed		
	progressive	pulsed on day 7 with			tumors, and demonstrated		
	metastatic	KLH and one of three			phenotypic characteristics of		
	BrCa.	peptides (hTERT I540,			tumor-lytic CD8+ T-cells		
		HIV RT-pol476 or					
		influenza MP58)					
		* Eligible pts were s.c.					
		administered a total of					
		15×10^6 autologous DCs					
		every other week for up					
		to six Vx					
Svane-2004	HLA-A2+ pts	* DC peptide p53	no maturation	6	* Specific T-cell responses	* SD was seen in 2 of	* Vx were well
[23]/ Phase	with	vaccine: autologous			against modified and	6 pts, 1 pt had a	tolerated
I	progressive	dendritic cells (DCs)			unmodified p53 peptides	transient regression	* No irritation
	advanced	loaded with 3 wild-type			observed in 3 pts, including 2 of	of a single lymph	at the site of
	BrCa.	and 3 P2 anchor			the pts with a possible clinical	node, and 1 had a	injection; no
		modified HLA-A2			benefit from the treatment.	mixed response.	allergic or
		binding p53 peptides					autoimmune
		(designed to increase					reactions; no
		HLA-A*0201 binding					haematologic,

		capacity and induction					hepatic,
		of p53-specific cytotoxic					pulmonary or
		T lymphocytes)					renal toxicities
		* PBMCs cultured for 7					* Most
		days with IL-4 and GM-					common side
		CSF and then frozen					effect: mild to
		* Cells pulsed with six					moderate local
		HLA-A2-associated p53					reaction at the
		peptides, and a pan-					site of
		MHC class II peptide,					Proleukin
		PADRE, for 2 h at 37C.					injection. All
		* Pts received up to 10					pts
		s.c. Vx with 5×10^6 p53-					experienced
		peptide loaded DC with					mild flu-like
		1–2 weeks interval.					symptoms
		Concomitantly, IL-2					lasting 12–24 h
		was administered s.c.					after injection
							of Proleukin.
Avigan-	Pts with	* DC vaccine: prepared	no maturation	23	* Fusion cells coexpressed	* 2 pts with breast	* No
2004 [40]/	metastatic	by fusing autologous		(10	tumor and DC antigens and	cancer showed	significant
Phase I	BrCa and	tumor cells and DCs		BrCa,	stimulated allogeneic T-cell	disease regressions,	treatment-
	renal cancer	* Tumor tissue was		13 RC)	proliferation.	including a near	related
	(RC), with	disrupted into single			* In a subset of pts, an increased	complete response of	toxicity
	tumor lesions	cell suspensions.			percentage of CD4 and CD8+ T-	a large chest wall	* No clinical
	accessible to	* PBMCs by			cells expressing intracellular	mass	evidence of
	biopsy or	leukapheresis were			IFN-gamma in response to in	* SD in 5 pts with RC	autoimmunity
	resection	cultured in GM-CSF, IL-				and 1 pt with BrCa	

							
		4, and autologous			vitro exposure to tumor lysate		
		plasma. Tumor cells			was observed		
		and DCs were					
		cocultured with PEG to					
		generate the fusions.					
		* Fusion cells were					
		administered s.c. Q3W					
Danet-	HLA-A2-	* DC vaccine:	no maturation	5	* No significant decline in the	See Vonderheide,	See
Desnoyers	positive pts	autologous DCs loaded			frequency of granulocyte,	2004	Vonderheide,
-2005 [24]/	with	with hTERT peptide			macrophage or erythroid CFCs		2004
Phase I	hormone-	and KLH			using CFC assays or long-term		
	independent	* From PBMCs. DCs			in vitro cultures		
	prostate	matured with GM-CSF			* In NOD/SCID mice, human		
	cancer or	and IL-4, and pulsed on			hematopoietic reconstitution		
	refractory	day 7 with KLH and			was easily detected, without		
	progressive	one of three peptides			quantitative or qualitative		
	metastatic	(hTERT I540, HIV RT-			differences between pre- and		
	BrCa.	pol476 or influenza			postvaccine samples		
		MP58).			•		
		* Eligible pts were s.c.					
		administered a total of					
		15 × 10 ⁶ autologous DCs					
		every other week for up					
		to six Vx.					
		IU SIX VX.					

Svane-2007	HLA-A2+ pts	* DC vaccine:	no maturation	26	* Therapy-induced p53 specific	* 19 pts available for	* Vaccine was
[25]/ Phase	with	autologous DCs loaded			T-cells were observed in 4/7 pts	first evaluation after	well tolerated.
II	progressive	with 3 wild-type and 3			with SD but only in 2/9 pts with	6 Vx	* No skin
	advanced	P2 anchor modified			PD.	* SD or minor	toxicity at the
	BrCa.	HLA-A2 binding p53				regression observed	site of vaccine
		peptides (designed to			* See Svane, 2008	in 8/19 evaluable pts	injection
		increase HLA-A*0201				or minor regression	* No
		binding capacity and				* PD in 11/19 pts	autoimmunity
		induction of p53-					* Most
		specific cytotoxic T					common side
		lymphocytes)					effect: mild to
		* PBMCs cultured for 7					moderate local
		days with IL-4 and GM-					reaction at the
		CSF and frozen					site of
		* Cells pulsed with six					proleukine
		HLA-A2-associated p53					injection
		peptides, and a pan-					* Pts
		MHC class II peptide,					experienced
		PADRE, for 2 h at 37°C					CTC grade 1–2
		* Pts received up to 10					flu-like
		s.c. Vx with 5×10^6 p53-					symptoms 12–
		peptide loaded DC with					24 h after
		1–2 weeks interval.					proleukine
		Concomitantly, IL-2					injection
		was administered s.c.					

Svane-2008	HLA-A2+ pts	* DC vaccine:	no maturation	26	* Any significant differences	See Svane, 2007	See Svane,
[26]/ Phase	with	autologous DCs loaded			between SD and PD pts		2007
II	progressive	with 3 wild-type and 3			* Decrease in naïve T-cells		
	advanced	P2 anchor modified			during vaccination, in		
	BrCa.	HLA-A2 binding p53			particular in pts with PD.		
		peptides (designed to			* The frequency of CD4+		
		increase HLA-A*0201			CD25high T-cells was almost		
		binding capacity and			doubled after only four weeks		
		induction of p53-			of weekly vaccination and IL-2		
		specific cytotoxic T			dosing.		
		lymphocytes).			* More than 90% of the CD4+		
		* PBMCs cultured for 7			CD25 high T-cells co-expressed		
		days with IL-4 and GM-			foxp3 confirming the		
		CSF and frozen.			regulatory functionality of		
		* Cells pulsed with six			these T-cells, independent of		
		HLA-A2-associated p53			clinical response		
		peptides, and a pan-					
		MHC class II peptide,					
		PADRE, for 2 h at 37°C.					
		* Pts received up to 10					
		s.c. Vx with 5×10^6 p53-					
		peptide loaded DC with					
		1–2 weeks interval.					
		Concomitantly, IL-2					
		was administered s.c.					

Santin-	pts with stage	* THER; DC protein	IL-1β, PGE2,	10	* All pts developed CD4(+) T-	* No sign of tumor	* Well
2008 [29]/	IB or IIA	HPV16/18 E7/KLH	TNF- <mark>α</mark>		cell and antibody responses to	recurrence detected	tolerated, no
Phase I	cervical	vaccine.			DC vaccination	in any of the treated	significant
	cancer (non	* PBMCs cultured with			* 8 out of 10 pts demonstrated	pts up to the time of	toxicities
	parametrial	GM-CSF			levels of E7-specific CD8(+) T-	writing	recorded
	involvement),	* At day 4, DCs were			cell counts		* No
	tumor HPV	pulsed overnight (12 to			* Vaccine dose did not predict		significant
	16/18+.	16 h) with HPV16/18 E7			the magnitude of the antibody		local or
		proteins and KLH at a			or T-cell response or the time to		systemic
		dose of 50 µg/ml			detection of HPV16/18 E7-		reactions
		* At day 5, maturation			specific immunity.		* No allergic
		was induced by 48h			* DTH responses to intradermal		reacions
		incubation with TNF-a,			injections of HPV E7 antigen		* No alteration
		IL-1b and PG-E2a			and KLH were detected for all		detected in
		* HPV16/18 E7/KLH-			pts after vaccination.		liver and renal
		pulsed DC were					function
		injected s.c. 10 cm					* Local
		inferior to the inguinal					reactions
		ligament of the anterior					detected (mild
		mid-thigh.					erythema,
		* Five DC Vx were					swelling/
		performed Q21D					induration,
							and pruritus)
							at the s.c.
							vaccination
							sites

Wang-2009	pts with stage	* THER; DC protein	IL-1β, PGE2,	8	* 12 T-cell lines from 8 subjects	See Santin, 2008.	See Santin,
[30]/	IB or IIA	HPV16/18 E7/KLH	TNF-α		(7 HPV 16-positive, 1 HPV 18-		2008.
Retrospect	cervical	vaccine.			positive) evaluated		
ive	cancer (non	* PBMCs cultured with			* Positive T-cell responses in 4		
	parametrial	GM-CSF			subjects (all HPV 16-positive),		
	involvement),	* At day 4, DCs were			all positive for the HPV 16 E7		
	tumor HPV	pulsed overnight (12 to			46-70		
	16/18+.	16 h) with HPV16/18 E7			(EPDRAHYNIVTFCCKCDSTL		
		proteins and KLH at a			RLCVQ) region		
		dose of 50 µg/ml			* T-cell clones specific for the		
		* At day 5, maturation			E7 47-70 region were isolated		
		was induced by 48h			from 1 subject		
		incubation with TNF-a,			* Further analyses revealed a		
		IL-1b and PG-E2a			novel, naturally processed, CD4		
		* HPV16/18 E7/KLH-			T-cell epitope, E7 58-68		
		pulsed DC were			(CCKCDSTLRLC), restricted by		
		injected s.c. 10 cm			the HLA-DR17 molecule		
		inferior to the inguinal					
		ligament of the anterior					
		mid-thigh.					
		* Five DC Vx were					
		performed Q21D					
Baek-2011	Advanced	* DC vaccine: DCs	IFN-γ	10	* Peripheral blood lymphocyte	* Clinical response	* Well
[32]/ Phase	RCC or BrCa.	pulsed with autologous		(6	proliferation and the number of	was observed in one	tolerated
I/II		tumor lysate (WTL) and		RCC,	IFN-r secreting cells were	RCC pt as SD	without major
		KLH.		4	induced in 6 pts without clear	* 9 cases showed PD	side effects
		* Cancer pts were		BrCa)	correlation with clinical		

		treated twice with			responses.		
		autologous CD34+			* NK activity was induced		
		hematopoietic stem cell-			significantly in 6 pts after		
		derived, GM-CSF/IFN-			vaccination.		
		γ -differentiated DCs			* DC vaccine-related decrease		
		(cultured during 7+7			of TGF- β level or increase of IL-		
		days), pulsed with			12p70 level and decline of		
		autologous WTL and			CD4+CD25+ T-cells were		
		KLH, Q4W.			observed in 3 pts		
		* Following each s.c					
		injection of therapeutic					
		DCs, low-dose (200					
		MIU) IL-2 was					
		introduced for 14					
		consecutive days					
Chu-2012	HLA-A2-	* DC vaccine: hTERT,	membrane	14	* Pts receiving Cy had a	* 3-year OS 90% (no	* No grade 3/4
[27]/ Phase	positive pts	Her2/neu, PADRE.	fractions from	OvCa	transient reduction in	difference for pts	vaccine-
I/II	with	* Monocyte-derived	Klebsiella		neutrophils, but no change in	receiving Cy over	related
	advanced	DCs loaded with	pneumoniae and		total lymphocytes or regulatory	controls)	toxicities were
	epithelial	synthetic peptides, with	IFN-γ		T-cells	* Estimated 3-year	noted
	OvCa or	or without low-dose			* Modest T-cell responses to	PFS was 40% versus	* Most
	primary	intravenous Cy. All pts			Her2/neu and hTERT were seen	80% for Arms 1	common
	peritoneal	also received			post-vaccine by IFN-γ ELISPOT	(without Cy) and 2	study-related
	cancer in	pneumococcal vaccine.			* Pts demonstrated below	(with cy),	toxicities:
	remission.	* PMBCs by			normal responses to the	respectively; the	reactogenicity
		leukapheresis cultured			diphtheria conjugate protein	estimated 3-year OS	as indicated
		in the presence of GM-				was 80 and 100%,	by erythema,

		CSF and IL-13 for 7			CRM197, a component of the	respectively	induration,
		days to generate DCs			pneumococcal vaccine	* Of 11 pts, 2 recurred	pruritus, and
		* Maturation by culture				during vaccination.	pain at the site
		with membrane				Nine received all 4	of injection,
		fractions from Klebsiella				doses: 3 pts recurred	fever and
		pneumoniae and IFN- γ				at 6, 17, and 26	fatigue
		* Mature DCs were				months, respectively,	
		pulsed with HLA-A2-				and 6 have no	
		restricted hTERT 988Y,				evidence of disease at	
		Her2/ neu 369V2V9,				36 months.	
		Her2/neu 689, and					
		PADRE peptides and					
		cryopreserved					
		* Each dose was injected					
		into the medial thighs at					
		24 intradermal sites					
Qi-2012	Double-	* DC vaccine:	IL-1β, PGE2,	31	* DC vaccines elicited Th1	* No difference in OS	* No
[33]/	negative	autologous dendritic	TNF-α		cytokine secretion and	between the pts with	unanticipated
Feasibility	stage II/IIIA	cells pulsed with			increased NK cells, CD8+ IFN-+	and without DC	or SAEs
	BrCa.	autologous tumor			cells but decreased the	vaccine	* A self-
		lysates (WTL)			percentage of CD3+ T-cells and	* 3-year PFS was	limited wheal-
		* DC vaccines generated			CD3+ HLA-DR+ T-cells in the	significantly	and-flare skin
		from CD14+ precursors			peripheral blood	prolonged: 76.9%	reaction
		pulsed with autologous			* Approximately 58% (18/31) of	versus 31.0% (with	appeared at
		WTL			pts had a DTH-positive	vs. without DC	the injection
		* DCs were matured			reaction	vaccine, $p < 0.05$).	site 24–48 h
		GM-CSF and IL-4					after

		* Tumor antigens added					immunization
		on day 5. On day 6, Il-					in 20 of 31
		1β, PGE2 and TNF- α					subjects
		added					during at least
		* Cells harvested on day					one of the four
		7 and directly used.					immunization
		Individuals were					s
		immunized					* No
		intradermally 4 times					abnormalities
							in
							hematological
							parameters or
							serum
							chemistries
Chiang-	Recurrent	* DC vaccine: OCDC	LPS, IFN-γ	5	* Subjects' DCs behaved similar	* 2 subjects remained	* All vaccines
2013 [34]/	OvCa.	* DC vaccine pulsed		OvCa	to normal donor DCs,	in remission for a	well tolerated;
Pilot		with HOCl-oxidized			producing high levels of IL-12	period of time much	most toxicities
		tumor lysate (WTL; see			and other important Th1	longer than expected	were <grade 2<="" td=""></grade>
		OvCa-Vac-Kandalaft-			cytokines and chemokines	based on historic	* Common
		2013).			* Following vaccination, a	observations	side effect: flu-
		* OCDC were			potent systemic inflammatory	* Both subjects	like
		administered through			activation was confirmed in	demonstrated a PFS2	symptomatolo
		direct injection into the			these subjects	> PFS1 in response to	gy (i.e. fatigue,
		one to two groin lymph			* Tumor-reactive T-cells	OCDC vaccination as	fever and
		nodes bilaterally under			exhibited a strong Th1	second-line therapy	chills)
		ultrasound guidance			polarization	following relapse	
					* OCDC vaccine was highly		

		* All subjects completed			efficient in crosspresentation to	from first-line	
		5 Vx (Figure 4A), except			CD8+ T-cells. This vaccine also	therapy	
		S1 who withdrew after			efficiently primed tumor-		
		3 Vx due to disease			specific CD4+ T-cell response.		
		progression.					
Kandalaft-	Relapsed	* DC vaccine: OCDC.	LPS, IFN-γ	25	* Day-4 DCs generated with	No data	No data
2013 [35]/	epithelial	* DC vaccine pulsed		OvCa	this protocol are similar to		
Phase I	carcinoma	with autologous			"classic" Day-7 DCs, in terms of		
	arising in the	oxidized WTL, in			phenotype and phagocytic		
	ovary,	combination with			capability, and have a higher		
	fallopian	antiangiogenesis			capacity than Day-7 DCs to		
	tube, or	therapy (bevacizumab)			produce IL-12p70 following		
	peritoneum	and metronomic Cy			LPS and IFN- γ stimulation		
		(three-arms study)			* In addition, these Day-4 DCs		
		* Faster, four-day			were highly immunogenic		
		protocol for DC					
		preparation, using GM-					
		CSF, IL-4 and serum-					
		free AIM-V media					
Kandalaft2	Recurrent	* DC vaccine: OCDC.	LPS, IFN-γ	6	* All subjects exhibited a	* 4 of 6 pts (66%)	* All vaccines
-2013 [36]/	OvCa pts for	* DC vaccine pulsed			dampened T-cell response to	achieved clinical	well tolerated
Pilot	whom tumor	with autologous			the diphtheria carrier protein	benefits with the	and no grade >
	lysate was	oxidized WTL, in			CRM197, given along with the	combination of	2 toxicities.
	available	combination with			first Vx to monitor immune	bevacizumab,	* Most
	from prior	antiangiogenesis			responsiveness	metronomic Cy and	frequent AE:
	cytoreductive	therapy (bevacizumab)			* DC Vx induced an immune	the vaccine (2 PR and	grade 1 or 2
	surgery.	and metronomic Cy			response against WTL and	2 SD)	hypertension

* 1 subject: post-(3 (three-arms study) specific immune responses * PBMCs cultured for 6 against peptides of known vaccine remision of occurrences), tumor-associated antigens such attributed to days with IL-4 and GM-14 mo in spite of a CSF. Immature DCs as HER2 prior PFS of 7 mo bevacizumab incubated for 18h with * Increased IgM seropositivity post-vaccine was detected the oWTL, IL-4 and **GM-CSF** * Frequency of tumor-specific * Pts underwent T-cells elicited by vaccine quite low (< 1 tumor reactive T-cell conditioning with per 500 PBLs) intravenous bevacizumab and oral metronomic Cy, sequentially followed by (1) bevacizumab plus vaccination with DCs pulsed with autologous tumor cell lysate supernatants, (2) lymphodepletion and (3) transfer of 5×10^9 autologous vaccineprimed T-cells in combination with the vaccine

Kobayashi	Recurrent	* DC vaccine: WT1,	OK-432, PG-E2	56	* No remarkable changes	* Clinical response	* Tolerable in
-2014 [28]/	OvCa, any	MUC1, CA125.		OvCa	observed in CD4+ T-cell, CD8+	evaluated in 56 pts at	all pts
Pilot	HLA type.	* Each pt was first			T-cell, and NK cell frequencies	3 and 6 months. The	* Most
		evaluated for human			after vaccination; in addition,	MST from diagnosis	common:
		leukocyte antigen			none of these factors affected	was 30.4 months and	injection site
		(HLA) expression, to			the median survival time (MST)	that from the first	reaction (68%)
		determine the type of			* No significant differences in	vaccination was 14.5	and fever
		peptides for			clinical outcomes between pts	months.	(32%)
		administration			with WT1-specific CTL increase	* The 1- and 2-y OS	* No serious
		* PBMCs from			and those without such an	from diagnosis were	acute allergic
		leukapheresis cultured			increase (17 pts evaluable)	87% and 65%, resp.	reaction such
		for 5 days in medium				At the time of the	as
		with GM-CSF and IL-4				final analysis, 35 pts	anaphylaxis,
		to generate immature				(63%) had died of	or other
		DCs				cancer.	common AEs
		* Maturation with OK-				* At 3 mo after the	such as
		432 and PG-E2 for 24h,				first Vx, any pt	arthralgia and
		then pulsed with the				showed CR.	elevated liver
		selected peptides, and				However, 2 pts	enzyme levels
		cryopreserved				(3.6%) had PR, 14	* No grade
		* All pts were i.d.				(25%) SD, 32 (57%)	3–4 toxicity or
		injected 5-7 times with				PD, and 8 (14%) were	evidence of
		DCs (approximately 107				not evaluated. The	autoimmune
		cells/injection) in close				DCR and ORR were	sequelae
		proximity to the axial				29% and 3.6%, resp.	
		and/or inguinal lymph					
		nodes. Injections were					

		repeated every 14–21					
		days					
		* OK-432, a					
		streptococcal					
		immunological					
		adjuvant, was					
		administered					
		simultaneously with the					
		DC vaccine to pts					
		without serious					
		allergies to penicillin or					
		other drugs					
Ramanath	Advanced,	* THER; DC WTL	IL-1 β , TNF- α	14	* One pt showed improved	* One pt who	* According to
an-2014	recurrent	vaccine.			proliferation of lymphocytes	received WTL-	WHO criteria,
[37]/ Phase	cervical	* PBMCs were cultured			after the third vaccination but	primed DCs and later	grade 0 or
I	cancer pts	for 7 days with GM-CSF			the increase however was not	cis-platin	grade one
	who had	and IL-4. Antigen			significant (paired two tailed t	chemotherapy	toxicity was
	failed	loading was evaluated			test =0.06)	showed a CR of her	observed in
	conventional	using three strategies:			* No significant proliferation	large metastatic	three pts
	therapy,	tumor lysate, cervical			responses were seen in any of	disease and remained	* None of the 9
	HPV+.	cancer cell line lysate			the other pts	disease free for more	pts evaluated
		(from HeLa, SiHa and			* Comparing arm II and arm III	than 72 months	had any
		C33A cells) and tumor			pts' responses also did not		elevation of
		RNA. Tumor lysate was			show significant difference		auto antibody
		added to immature DC.					levels after the
		* After 4 h of exposure					third
		to antigen, IL-1β and					vaccination

		TNF- α were added					although 1 pt,
		and the cells were					who was in
		incubated for 3 days.					arm I (saline
		On day 10, the mature					only) showed
		DC were frozen.					slight
		* Vx were given to each					elevation of
		pt intra-dermally. Pts					ANA levels
		received three Vx, one					
		every 14 days. A					
		maximum of 1x106 cells					
		were given at each					
		vaccination					
Baek-2015	OvCa pts	* DC aWTL vaccine:	TNF-α	10	* In the 3 pts with disease free	* In 3 out of 10 pts,	The
[38]/ Phase	with minimal	DCs pulsed with			long-term survival, significant	the inclusion status	vaccination
I/II	residual	autologous tumor lysate			immune alterations were	after the initial	was well
	disease	and KLH.			observed, including increased	therapy showed the	tolerated. The
		* Cancer pts received			natural killer (NK) activity,	maintenance of CR	most common
		GM-CSF (injected s.c.)			IFN-γ-secreting T-cells,	after DC vaccination	side effects
		during 5 days, for			immune-stimulatory cytokine	for 83, 80.9 and 38.2	were flu-like
		monocyte mobilization,			secretion and reduced immune-	months without	symptoms.
		and PBMCs were then			suppressive factor secretion	disease relapse	
		collected by			after DC vaccination	* 1 pt with SD	
		leukapheresis. Cells				experienced the	
		were cultured with CM-				complete	
		CSF and IL-4 for 7 days.				disappearance of	
		On day 7, TNF-a was				tumor after DC	
		added for 2 days				vaccination,	

		* DCs were pulsed with				maintained for 50.8	
		aWTL and KLH on day				months until tumor	
		8 O/N. On day 9, cells				recurrence	
		were harvested,				* In 2 pts with PR was	
		washed, resuspended				not responding to DC	
		and tested				vaccination and their	
		* Three to seven months				disease recurred	
		after the initial surgery					
		and chemotherapy, 10					
		pts were treated with					
		aWTL-loaded DCs					
		(4.13×10 ⁷ ±0.27×10 ⁷					
		cells/injection), followed					
		by 14 consecutive IL-2					
		(200 mIU) injections in a					
		single vaccination					
		protocol					
		* Two Vx were s.c.					
		administered in an area					
		adjacent to the axillary					
		lymph node Q4W					
Tanyi-2018	platinum-	* Pilot clinical trial	LPS, IFN-γ	6	* Vx induced T-cell responses		* A total of 39
[39]/ Pilot	treated,	testing a personalized			to autologous tumor antigen		vaccine doses
	immunothera	vaccine (OCDC)			associated with significantly		were
	py-naïve,	generated by			prolonged survival		administered
	recurrent	autologous DCs pulsed			* Vx also amplified T-cell		without
	OvCa pts	with oxidized			responses against mutated		

autologous whole-	neoepitopes derived from	serious
tumor cell lysate	nonsynonymous somatic tumor	adverse events
(aWTL), which was	mutations, including priming	
injected intranodally in	of T-cells against previously	
platinum-treated,	unrecognized neoepitopes, as	
immunotherapy-naïve,	well as novel T-cell clones of	
recurrent ovarian cancer	markedly higher avidity	
pts	against previously recognized	
	neoepitopes	

Abbreviations: aWTL, autologous whole tumor lysate; BrCa, breast cancer; CR, complete response; Cy, cyclophosphamide; DC(s), dendritic cell(s)s; i.d., intradermal; i.v., intravenous; Matur, maturation; MEL, melanoma; OS: overall survival; OvCa, ovarian cancer; PFS, progression-free survival; ProstCa, prostate cancer; PR, partial response; Pt(s), patient(s); RC, renal cell carcinoma; resp., respectively; s.c., subcutaneous; SD, stable disease; UtSar, uterine sarcoma; Vx, vaccinations.