Clinical trial results:
An open Phase II study to assess the clinical activity and safety of recMAGE-A3 + AS15 cancer immunotherapeutic in patients with metastatic cutaneous melanoma, and to explore its immunogenic properties, including their relation to tumor infiltration, genomics and proteomics

Summary

EudraCT number	2008-001301-42
Trial protocol	FR BE
Global end of trial date	03 November 2014

Results information

Result version number	v2 (current)
This version publication date	28 April 2021
First version publication date	03 March 2016
Version creation reason	• Correction of full data set Results have been amended to account for consistency with other registries.

Trial information

Trial identification

Sponsor protocol code	111473

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT00896480
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	GlaxoSmithKline Biologicals
Sponsor organisation address	Rue de l'Institut 89, Rixensart, Belgium, B-1330
Public contact	Clinical Trials Call Center, GlaxoSmithKline Biologicals, 044 2089-904466, GSKClinicalSupportHD@gsk.com
Scientific contact	Clinical Trials Call Center, GlaxoSmithKline Biologicals, 044 2089-904466, GSKClinicalSupportHD@gsk.com

Notes:

Paediatric regulatory details

Is trial part of an agreed paediatric investigation plan (PIP)	No
Does article 45 of REGULATION (EC) No 1901/2006 apply to this trial?	No
Does article 46 of REGULATION (EC) No 1901/2006 apply to this trial?	No

Notes:

Results analysis stage

Analysis stage	Final
Date of interim/final analysis	06 August 2015
Is this the analysis of the primary completion data?	Yes
Primary completion date	03 November 2014
Global end of trial reached?	Yes
Global end of trial date	03 November 2014
Was the trial ended prematurely?	Yes

Notes:

General information about the trial

Main objective of the trial:

Clinical Activity - To characterize in patients with MAGE-A3-positive metastatic cutaneous melanoma:
The clinical activity of the MAGE-A3 ASCI study treatment in terms of objective response (OR), stable disease (SD) and mixed response (MR)*
The clinical activity of the MAGE-A3 ASCI study treatment in terms of time to treatment failure (TTF)*
The safety of the MAGE-A3 ASCI study treatment.
Immunogenicity - To document the humoral and cellular immune response induced by the MAGE-A3 ASCI study treatment.

Protection of trial subjects:

The patients will be observed closely for at least 30 minutes following the administration of treatments, with appropriate medical treatment readily available in case of a rare anaphylactic reaction. VMAGE-A3 ASCI/placebo were administered by qualified and trained personnel, only to eligible subjects with no contraindications to any components of these products. During treatment, the following was checked to assess need to postpone treatment: acute disease at time of administration; any systemic grade ≥ 2 Common Terminology Criteria Adverse Event related or possibly related to treatment; fever, defined as an oral, axillary or tympanic temperature =< 38°C; need for influenza vaccine, immunoglobulins and/or any blood products; any medical reason exposing the patient to unacceptable risk. Patients were required to discontinue treatment in case of evidence of disease relapse/occurrence of second primary lung cancer; treatment with either investigational or non-registered product other than MAGE-A3 ASCI study product or other anticancer treatments; anaphylactic reaction following treatment administration; any intolerable adverse event; clinical signs or symptoms indicative of any autoimmune disorder, except vitiligo; appearance of any confirmed or suspected immunosuppressive or immunodeficient condition, or any condition requiring use of any immunosuppressive agent or systemic corticosteroids prescribed for chronic use; inability of the patient to complete study evaluations due to unforeseen circumstances; other conditions indicating the patient's best interest to be withdrawn from treatment. In addition, between the end of the 120-weeks treatment phase, the following follow-up (FU) of patients was also planned: 1) an active FU for survival, recurrence, serious adverse events related to treatment & SAEs related to study participation and concurrent GSK medication of up to 5 years from the 1st treatment, and 2) annual contacts up to 10 years after 1st treatment.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	19 May 2009
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Belgium: 9
Country: Number of subjects enrolled	France: 15

Worldwide total number of subjects	24
EEA total number of subjects	24

Notes:

Subjects enrolled per age group	
In utero	0
Preterm newborn - gestational age < 37 wk	0
Newborns (0-27 days)	0
Infants and toddlers (28 days-23 months)	0
Children (2-11 years)	0
Adolescents (12-17 years)	0
Adults (18-64 years)	11
From 65 to 84 years	13
85 years and over	0

Subject disposition

Recruitment

Recruitment details: -

Pre-assignment

Screening details:

During the screening the following steps occurred: check for inclusion criteria, contraindications/precautions, medical history of the subjects and signing informed consent forms.

Period 1

Period 1 title	Overall trial (overall period)
Is this the baseline period?	Yes
Allocation method	Not applicable
Blinding used	Not blinded

Arms

Arm title	MAGE-A3 Group

Arm description:

Patients planned to receive intramuscularly up to 24 doses of MAGE-A3 ASCI (the study product), in 4 cycles.

Arm type	Experimental
Investigational medicinal product name	recMAGE-A3 recombinant protein formulated in AS15 adjuvant
Investigational medicinal product code	GSK2132231A
Other name	
Pharmaceutical forms	Powder and suspension for suspension for injection
Routes of administration	Intramuscular use

Dosage and administration details:

Administration as follows:
-Cycle 1 (ending Week 11): 6 doses at 2-week intervals (Weeks 1, 3, 5, 7, 9 and11)
-Cycle 2 (ending Week 30): 6 doses at 3-week intervals (Weeks 15, 18, 21, 24, 27 and 30)
-Cycle 3 (ending Week 52): 4 doses at 6-week intervals (Weeks 34, 40, 46 and 52)
-Cycle 4: 4 doses at 12-week intervals, starting 12 weeks after end of Cycle 3, followed by, after an interruption of treatment of 6 months, 4 doses at 24-week intervals. All analyses were performed on the overall study population (MAGE3 Group) as well as in the subsets of patients with or without the pre-specified gene signature (GS+ or GS- groups) and in one patient with unknown status as regards GS signature (Unknown Group).

Number of subjects in period 1	MAGE-A3 Group
Started	24
Completed	3
Not completed	21
Other Disease progression/recurrence	21

Baseline characteristics

Reporting groups

Reporting group title	MAGE-A3 Group

Reporting group description:

Patients planned to receive intramuscularly up to 24 doses of MAGE-A3 ASCI (the study product), in 4 cycles.

Reporting group values	MAGE-A3 Group	Total	
Number of subjects	24	24	
Age categorical Units: Subjects			
In utero		0	
Preterm newborn infants (gestational age < 37 wks)		0	
Newborns (0-27 days)		0	
Infants and toddlers (28 days-23 months)		0	
Children (2-11 years)		0	
Adolescents (12-17 years)		0	
Adults (18-64 years)		0	
From 65-84 years		0	
85 years and over		0	
Age continuous Units: years			
arithmetic mean	65.4		
standard deviation	± 12.4	-	
Gender categorical Units: Subjects			
Female	17	17	
Male	7	7	

Subject analysis sets

Subject analysis set title	GS+/+ Group
Subject analysis set type	Sub-group analysis

Subject analysis set description:

Subset of patients with the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature positive for two biopsies, as assessed at screening.

Subject analysis set title	GS+/- Group
Subject analysis set type	Sub-group analysis

Subject analysis set description:

Subset of patients the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature positive for only one biopsies, as assessed at screening.

Subject analysis set title	GS- Group
Subject analysis set type	Sub-group analysis

Subject analysis set description:

Subset of patients the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature negative for both biopsies, as

assessed at screening.

Reporting group values	GS+/+ Group	GS+/- Group	GS- Group
Number of subjects	8	8	8
Age categorical Units: Subjects			
In utero			
Preterm newborn infants (gestational age < 37 wks)			
Newborns (0-27 days)			
Infants and toddlers (28 days-23 months)			
Children (2-11 years)			
Adolescents (12-17 years)			
Adults (18-64 years)			
From 65-84 years			
85 years and over			
Age continuous Units: years			
arithmetic mean	70.1	66.3	59.9
standard deviation	± 13.4	± 12.7	± 10.1
Gender categorical Units: Subjects			
Female	5	5	7
Male	3	3	1

End points

End points reporting groups

Reporting group title	MAGE-A3 Group
Reporting group description:	
Patients planned to receive intramuscularly up to 24 doses of MAGE-A3 ASCI (the study product), in 4 cycles.	
Subject analysis set title	GS+/+ Group
Subject analysis set type	Sub-group analysis
Subject analysis set description:	
Subset of patients with the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature positive for two biopsies, as assessed at screening.	
Subject analysis set title	GS+/- Group
Subject analysis set type	Sub-group analysis
Subject analysis set description:	
Subset of patients the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature positive for only one biopsies, as assessed at screening.	
Subject analysis set title	GS- Group
Subject analysis set type	Sub-group analysis
Subject analysis set description:	
Subset of patients the pre-specified gene signature, receiving the MAGE-A3 ASCI product. Gene-signature sub-grouping was based on patients having a gene signature negative for both biopsies, as assessed at screening.	

Primary: Number of patients with mixed response (MxR) to MAGE-A3 ASCI study treatment

End point title	Number of patients with mixed response (MxR) to MAGE-A3 ASCI study treatment[1]
End point description:	
Assessment was done based on a set of measurable lesions(MLs) identified at baseline as target lesions(TLs) and non-target lesions(NTLs) followed up until disease progression.MLs were assessed as regards matching below MR definitions. If Evaluability per RECIST: a)MR Type 1= at least (a.l.) 30% decrease in longest diameter(LD) in a.l. 1 TL measured at baseline.Such response occurring in SD/PD status of LD of TL and without appearance of 1 or more new lesions (= SD/PD with TL regression); b)MR Type 2: appearance of 1 or more new lesions occurring in SD/PR status of LD of TL(= SD/PR with new lesion).If Non-evaluability per RECIST (due to LD<20mm): a)MR Type 1 = clear decrease in diameters occurring in a.l. 1 TL measured at baseline.Such response occurring in SD/PD status of LD of (baseline) TL and without appearance of 1 or more new lesions (= SD/PD with TL regression); b)MR Type 2 = appearance of 1 or more new lesions occurring in SD/PR status of LD of TL (= SD/PR with new lesion)	
End point type	Primary
End point timeframe:	
From Pre-treatment (up to 4 weeks before first treatment) to study end (Year 4), each patient being censored out of the analysis at 1st report of disease progression in assessed lesions	

Notes:

[1] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group	GS+/+ Group	GS+/- Group	GS- Group
Subject group type	Reporting group	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	24	8	8	8
Units: Patients				
Best response CR	2	2	0	0
Best response PR	2	1	1	0
Best response MR (SD/PR)	2	1	1	0
Best response MR (SD/PD)	3	2	0	1
Best response SD	1	0	0	1
Best response SD/PD	0	0	0	0
Best response PD (SPD)	4	1	1	2
Best response PD (SPD/MR)	10	1	5	4
Best response NE	0	0	0	0

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with best objective tumor response (OR) to MAGE-A3 ASCI study treatment

End point title	Number of patients with best objective tumor response (OR) to MAGE-A3 ASCI study treatment[2]

End point description:

Response assessment was done based on a set of MLs identified at baseline as TLs, and followed up until disease progression. OR was defined as the best Overall Response (OR) in a patient. OR = Complete Response (CR) + Partial Response (PR). Responses were categorized as CR, PR, stable disease (SD), SD/PR, progressive disease (PD) and non-evaluable (NE). Per Response Evaluation Criteria In Solid Tumors Criteria (RECIST v1.0) for target lesions and assessed by Magnetic-resonance imaging: Complete Response (CR) = disappearance of all target lesions; Partial Response (PR) = ≥30% decrease in the sum of the longest diameter of target lesions; Stable Disease (SD) = neither sufficient shrinkage to be PR, not sufficient increase to qualify for Progressive Disease (PD); PD = ≥20% increase in the sum of largest diameter for target lesions. Best objective response = PR or CR. Disease control = CR, or PR, or SD, or SD/PR.

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to study end (Year 4), each patient being censored out of the analysis at 1st report of disease progression in assessed lesions

Notes:

[2] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group	GS+/+ Group	GS+/- Group	GS- Group
Subject group type	Reporting group	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	24	8	8	8
Units: Patients				
Best response CR	2	2	0	0
Best response PR	2	1	1	0
Best response SD	2	0	1	1
Best response SD/PR	1	1	0	0
Best response PD	17	4	6	7

Best response NE	0	0	0	0
Best objective response Yes	4	3	1	0
Best objective response No	20	5	7	8
Disease Control Yes	7	4	2	1
Disease Control No	17	4	6	7

Statistical analyses

No statistical analyses for this end point

Primary: Time to treatment failure (TTF), by Gene Signature

End point title	Time to treatment failure (TTF), by Gene Signature[3]

End point description:

TTF was defined as withdrawal from treatment with the MAGE-A3 ASCI study product due to disease progression or death. TTF analysis was performed using the non-parametric Kaplan-Meier method. "9999" as placeholder value for confidence interval result being not applicable/missing: upper limit not calculated as 1 patient (among 8) still under treatment at the time of the analysis.

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to study end (Year 4), each patient being censored out of the analysis at 1st report of disease progression in assessed lesions

Notes:

[3] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	GS+/+ Group	GS+/- Group	GS- Group	
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	
Number of subjects analysed	8	8	8	
Units: Months				
median (confidence interval 95%)				
TTF	14.8 (2.3 to 9999)	2.3 (0.5 to 15)	2.4 (0.5 to 4.6)	

Statistical analyses

No statistical analyses for this end point

Primary: Number of seroconverted patients for anti-MAGE-A3

End point title	Number of seroconverted patients for anti-MAGE-A3[4]

End point description:

Seroconversion was defined as a concentration of antibodies assessed that was greater than the cut-off value for a patient whose concentration of such antibodies was below the cut-off level before the initiation of treatment. Seroconverted patients were those patients with anti-MAGE-A3 antibody concentrations ≥ 27 EL.U/mL.

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients

withdrawn from study treatment before completion)

Notes:

[4] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group	GS+/+ Group	GS+/- Group	GS- Group
Subject group type	Reporting group	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	18	7	5	6
Units: Patients				
Anti-MAGE-A3, PRE [N=18;7;5;6]	3	1	1	1
Anti-MAGE-A3, D2 [N=17;7;4;6]	3	1	1	1
Anti-MAGE-A3, D7 [N=18;7;5;6]	4	1	2	1
Anti-MAGE-A3, D15 [N=17;7;5;5]	9	3	3	3
Anti-MAGE-A3, D16 [N=15;7;3;5]	8	3	2	3
Anti-MAGE-A3, W5 [N=16;7;4;5]	16	7	4	5
Anti-MAGE-A3, W11 [N=12;4;4;4]	12	4	4	4
Anti-MAGE-A3, W13 [N=16;7;4;5]	16	7	4	5
Anti-MAGE-A3, W21 [N=10;7;2;1]	10	7	2	1
Anti-MAGE-A3, W32 [N=10;7;2;1]	10	7	2	1
Anti-MAGE-A3, W54 [N=5;4;1;0]	5	4	1	0
Anti-MAGE-A3, W76 [N=3;3;0;0]	3	3	0	0
Anti-MAGE-A3, W78 [N=2;1;1;0]	2	1	1	0
Anti-MAGE-A3, W100 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W102 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W124 [N=2;1;1;0]	2	1	1	0
Anti-MAGE-A3, W126 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W148 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W150 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W174 [N=2;2;0;0]	2	2	0	0
Anti-MAGE-A3, CCL [N=5;2;0;3]	5	2	0	3

Statistical analyses

No statistical analyses for this end point

Primary: Anti-MAGE-A3 antibody concentrations

End point title	Anti-MAGE-A3 antibody concentrations[5]

End point description:

Anti-MAGE-A3 antibody concentrations were presented as geometric mean concentrations (GMCs) and expressed in ELISA units per millilitre (EL.U/mL). A seropositive patient was defined as a patient whose anti-MAGE-A3 antibody concentration greater than or equal to 27 EL.U/mL. D=Day, W=Week. Limits of the CI were entered = to the GMC when not available: limits are not available, as there is only one subject analyzed in the group, at this time point.

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion)

Notes:

[5] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group	GS+/+ Group	GS+/- Group	GS- Group
Subject group type	Reporting group	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	18	7	5	6
Units: EL.U/mL				
geometric mean (confidence interval 95%)				
Anti-MAGE-A3, PRE [N=18;7;5;6]	14.2 (9.5 to 21.4)	13.9 (6.2 to 31.4)	15.3 (4.7 to 49.9)	13.8 (6.1 to 31.3)
Anti-MAGE-A3, D2 [N=17;7;4;6]	14.6 (9.5 to 22.6)	13.9 (6.2 to 30.8)	16.8 (3.2 to 86.8)	14.2 (5.8 to 34.7)
Anti-MAGE-A3, D7 [N=18;7;5;6]	16.6 (11.1 to 24.7)	16.1 (6.9 to 37.4)	18.7 (5.9 to 59.2)	15.5 (7.3 to 32.7)
Anti-MAGE-A3, D15 [N=17;7;5;5]	56.6 (24.8 to 129)	48.5 (11.6 to 202.6)	72 (7.4 to 699.2)	55.1 (5.9 to 512.2)
Anti-MAGE-A3, D16 [N=15;7;3;5]	65.1 (25.5 to 166.6)	55.4 (11.8 to 260.6)	102.4 (0.7 to 15279.4)	62.2 (5.8 to 662.9)
Anti-MAGE-A3, W5 [N=16;7;4;5]	1865.7 (906.6 to 3839.3)	1396.9 (234.5 to 8321.5)	4120.2 (2093 to 8111)	1484.1 (736.8 to 2989.6)
Anti-MAGE-A3, W11 [N=12;4;4;4]	6312.1 (4177.9 to 9536.4)	4161.6 (1542.3 to 11229.3)	9775.9 (3065.3 to 31177.1)	6181.6 (3414.2 to 11191.8)
Anti-MAGE-A3, W13 [N=16;7;4;5]	9080.5 (7030.9 to 11727.6)	7797.4 (4564.5 to 13320.2)	13789.3 (7278.1 to 26125.3)	8046.1 (7249.1 to 8930.8)
Anti-MAGE-A3, W21 [N=10;7;2;1]	8540.2 (5768.3 to 12644)	7400.3 (4304 to 12724.1)	12544.5 (371.2 to 423881.3)	10790 (10790 to 10790)
Anti-MAGE-A3, W32 [N=10;7;2;1]	6826.9 (4728.9 to 9855.8)	6128 (3601.6 to 10426.5)	8384.1 (459.4 to 153006.6)	9641 (9641 to 9641)
Anti-MAGE-A3, W54 [N=5;4;1;0]	7429.9 (4876.9 to 11319.3)	6542.6 (4660.3 to 9185.1)	12357 (12357 to 12357)	0 (0 to 0)
Anti-MAGE-A3, W76 [N=3;3;0;0]	3539.3 (2578.1 to 4859)	3539.3 (2578.1 to 4859)	0 (0 to 0)	0 (0 to 0)
Anti-MAGE-A3, W78 [N=2;1;1;0]	5972.3 (24 to 1485256)	3869 (3869 to 3869)	9219 (9219 to 9219)	0 (0 to 0)
Anti-MAGE-A3, W100 [N=3;2;1;0]	5512.4 (1673 to 18163)	4260 (452.7 to 40086)	9230 (9230 to 9230)	0 (0 to 0)
Anti-MAGE-A3, W102 [N=3;2;1;0]	6523.7 (1713.7 to 24834.3)	4804.1 (1467.6 to 15725.7)	12030 (12030 to 12030)	0 (0 to 0)
Anti-MAGE-A3, W124 [N=2;1;1;0]	3429.1 (0.2 to 67399712)	1575 (1575 to 1575)	7466 (7466 to 7466)	0 (0 to 0)
Anti-MAGE-A3, W126 [N=3;2;1;0]	5459.7 (1022.8 to 29145.2)	3699.8 (2894.3 to 4729.5)	11889 (11889 to 11889)	0 (0 to 0)
Anti-MAGE-A3, W148 [N=3;2;1;0]	3010.6 (288.7 to 31395.3)	1754.4 (351.7 to 8751.9)	8865 (8865 to 8865)	0 (0 to 0)
Anti-MAGE-A3, W150 [N=3;2;1;0]	4657.1 (529.4 to 40967)	3077.8 (5.3 to 1801276)	10663 (10663 to 10663)	0 (0 to 0)
Anti-MAGE-A3, W174 [N=2;2;0;0]	2248.7 (226.4 to 22332.9)	2248.7 (226.4 to 22332.9)	0 (0 to 0)	0 (0 to 0)

Anti-MAGE-A3, CCL [N=5;2;0;3]	6972.9 (2571.8 to 18905.3)	4057.1 (0.1 to 135760000)	0 (0 to 0)	10004.8 (4092.9 to 24455.9)

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with treatment response for anti-MAGE-A3 antibodies

End point title	Number of patients with treatment response for anti-MAGE-A3 antibodies[6]

End point description:

For initially seronegative patients: post-administration antibody concentration ≥ 27 EL.U/mL
For initially seropositive patients: post-administration antibody concentration ≥ 2 fold the pre-vaccination antibody concentration

End point type	Primary

End point timeframe:

From Day 2 to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion)

Notes:

[6] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group	GS+/+ Group	GS+/- Group	GS- Group
Subject group type	Reporting group	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	18	7	5	6
Units: Patients				
Anti-MAGE-A3, D2 [N=17;7;4;6]	0	0	0	0
Anti-MAGE-A3, D7 [N=18;7;5;6]	1	0	1	0
Anti-MAGE-A3, D15 [N=17;7;5;5]	9	3	3	3
Anti-MAGE-A3, D16 [N=15;7;3;5]	8	3	2	3
Anti-MAGE-A3, W5 [N=16;7;4;5]	16	7	4	5
Anti-MAGE-A3, W11 [N=12;4;4;4]	12	4	4	4
Anti-MAGE-A3, W13 [N=16;7;4;5]	16	7	4	5
Anti-MAGE-A3, W21 [N=10;7;2;1]	10	7	2	1
Anti-MAGE-A3, W32 [N=10;7;2;1]	10	7	2	1
Anti-MAGE-A3, W54 [N=5;4;1;0]	5	4	1	0
Anti-MAGE-A3, W76 [N=3;3;0;0]	3	3	0	0
Anti-MAGE-A3, W78 [N=2;1;1;0]	2	1	1	0
Anti-MAGE-A3, W100 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W102 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W124 [N=2;1;1;0]	2	1	1	0
Anti-MAGE-A3, W126 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W148 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W150 [N=3;2;1;0]	3	2	1	0
Anti-MAGE-A3, W174 [N=2;2;0;0]	2	2	0	0
Anti-MAGE-A3, CCL [N=5;2;0;3]	5	2	0	3

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with CD4+ and CD8+ T cell frequency ≥ 1.24 cut-off

End point title	Number of patients with CD4+ and CD8+ T cell frequency ≥ 1.24 cut-off[7]

End point description:

A patient was considered as a cellular mediated immune (CMI) responder if there was an increased amount of antigen-specific T-cells after immunization as compared to the patient's baseline value. These specific T-cells included the CD4+ or CD8+ T-cells producing cytokines Tumor Necrosis Factor-alpha (TNF-α) and/or Interferon-gamma (INF-γ).

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion)

Notes:

[7] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: Patients				
CD4.TNFα (+) + IFNγ (+) PRE (N=17)	1			
CD8.TNFα (+) + IFNγ (+) PRE (N=18)	1			
CD4.TNFα (+) + IFNγ (+) (W5) (N=17)	15			
CD8.TNFα (+) + IFNγ (+) (W5) (N=17)	0			
CD4.TNFα (+) + IFNγ (+) (W13) (N=14)	11			
CD8.TNFα (+) + IFNγ (+) (W13) (N=15)	0			
CD4.TNFα (+) + IFNγ (+) (W32) (N=10)	9			
CD8.TNFα (+) + IFNγ (+) (W32) (N=10)	2			
CD4.TNFα (+) + IFNγ (+) (W54) (N=5)	4			
CD8.TNFα (+) + IFNγ (+) (W54) (N=5)	0			
CD4.TNFα (+) + IFNγ (+) (W78) (N=2)	2			
CD8.TNFα (+) + IFNγ (+) (W78) (N=2)	0			
CD4.TNFα (+) + IFNγ (+) (W102) (N=3)	2			
CD8.TNFα (+) + IFNγ (+) (W102) (N=3)	1			
CD4.TNFα (+) + IFNγ (+) CCL (N=2)	1			
CD8.TNFα (+) + IFNγ (+) CCL (N=2)	0			

CD4.TNFα (+) + IFNγ (+) At any time point (N=17)	15			
CD8.TNFα (+) + IFNγ (+) At any time point (N=17)	3			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with a cellular response (Anti-MAGE-A3 specific CD4+ and CD8+ T-cells concentrations after immunization)

End point title	Number of patients with a cellular response (Anti-MAGE-A3 specific CD4+ and CD8+ T-cells concentrations after immunization)[8]

End point description:

A patient was considered as a cellular mediated immune (CMI) responder if there was an increased amount of antigen-specific T-cells after immunization as compared to the patient's baseline value. These specific T-cells included the CD4+ or CD8+ T-cells producing cytokines Tumor Necrosis Factor-alpha (TNF-α) and/or Interferon-gamma (INF-γ).

End point type	Primary

End point timeframe:

From Week 5 to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion)

Notes:

[8] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	17			
Units: Patients				
CD4.TNFα (+) + IFNγ (+) (W5) (N=16)	5			
CD8.TNFα (+) + IFNγ (+) (W5) (N=17)	0			
CD4.TNFα (+) + IFNγ (+) (W13) (N=13)	8			
CD8.TNFα (+) + IFNγ (+) (W13) (N=15)	0			
CD4.TNFα (+) + IFNγ (+) (W32) (N=9)	4			
CD8.TNFα (+) + IFNγ (+) (W32) (N=10)	0			
CD4.TNFα (+) + IFNγ (+) (W54) (N=4)	2			
CD8.TNFα (+) + IFNγ (+) (W54) (N=5)	0			
CD4.TNFα (+) + IFNγ (+) (W78) (N=2)	0			
CD8.TNFα (+) + IFNγ (+) (W78) (N=2)	0			
CD4.TNFα (+) + IFNγ (+) (W102) (N=3)	1			
CD8.TNFα (+) + IFNγ (+) (W102) (N=3)	0			
CD4.TNFα (+) + IFNγ (+) CCL (N=2)	1			
CD8.TNFα (+) + IFNγ (+) CCL (N=2)	0			
CD4.TNFα (+) + IFNγ (+) At any time point (N=16)	12			

CD8.TNFα (+) + IFNγ (+) At any time point (N=17)	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients reported with ASCI-related grade3/4 adverse events (AEs) according to the Common Terminology Criteria (CTCAE) version 3.0.

End point title	Number of patients reported with ASCI-related grade3/4 adverse events (AEs) according to the Common Terminology Criteria (CTCAE) version 3.0.[9]

End point description:

The assessed AEs were ASCI-related grade 3/4 adverse events according to the Common Terminology Criteria for Adverse Events (CTCAE) version 3.0. An unsolicited AE covers any untoward medical occurrence in a clinical investigation patient temporally associated with the use of a medicinal product, whether or not considered related to the medicinal product. An AE can therefore be any unfavorable and unintended sign (including an abnormal laboratory finding), symptom, or disease (new or exacerbated) temporally associated with the use of a medicinal product. For marketed medicinal products, this also includes failure to produce expected benefits (i.e. lack of efficacy), abuse or misuse. AEs may include pre- or post-treatment events that occur as a result of protocol-mandated procedures (i.e. invasive procedures, modification of patient's previous therapeutic regimen).

End point type	Primary

End point timeframe:

Within the 31-day (Days 0-30) post-administration periods.

Notes:

[9] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Any event, Grade 3	0			
Any event, Grade 4	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients reported with serious adverse events (SAEs)

End point title	Number of patients reported with serious adverse events (SAEs)[10]

End point description:

Serious adverse events (SAEs) include medical occurrences that result in death, are life threatening, require hospitalization or prolongation of hospitalization or result in disability/incapacity, is a congenital anomaly/birth defect in the offspring of a patient, is a Grade 4 AE according to the CTCAE, version3.0. Events which were part of the natural course of the disease under study were captured as part of the clinical activity outcome variables in this study; therefore did not need to be reported as SAEs.

Progression/recurrence of the tumor was recorded as part of the clinical assessment data collection, and deaths due to progressive disease was recorded on a specific form, but not as an SAE. However, if the investigator considered that there was a causal relationship between treatment or protocol design/procedures and the disease progression/recurrence, then the event was reported as an SAE. Any new primary cancer (non-related to the cancer under study) was reported as an SAE.

End point type	Primary

End point timeframe:

During the entire study period (from Day 0 to CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion

Notes:

[10] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Any SAEs	2			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Alanine aminotransferase (ALT) values by maximum grade

End point title	Number of patients with abnormal Alanine aminotransferase (ALT) values by maximum grade[11]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality(UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. ALT – SCR G0; SE G3 = ALT with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[11] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
ALT - SCR G0; SE G0	20			
ALT - SCR G0; SE G1	3			
ALT - SCR G0; SE G2	0			

ALT - SCR G0; SE G3	1			
ALT - SCR G0; SE G4	0			
ALT - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Aspartate aminotransferase (AST) values by maximum grade

End point title	Number of patients with abnormal Aspartate aminotransferase (AST) values by maximum grade[12]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. AST – SCR G0; SE G3 = AST with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[12] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
AST - SCR G0; SE G0	19			
AST - SCR G0; SE G1	5			
AST - SCR G0; SE G2	0			
AST - SCR G0; SE G3	0			
AST - SCR G0; SE G4	0			
AST - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Alkaline Phosphatase (ALK) values by maximum grade

End point title	Number of patients with abnormal Alkaline Phosphatase (ALK) values by maximum grade[13]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria

Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. ALK – SCR G0; SE G3 = ALK with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[13] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
ALK - SCR UNK; SE G0	0			
ALK - SCR UNK; SE G1	0			
ALK - SCR UNK; SE G2	0			
ALK - SCR UNK; SE G3	0			
ALK - SCR UNK; SE G4	0			
ALK - SCR UNK; SE UNK	1			
ALK - SCR G0; SE G0	17			
ALK - SCR G0; SE G1	6			
ALK - SCR G0; SE G2	0			
ALK - SCR G0; SE G3	0			
ALK - SCR G0; SE G4	0			
ALK - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Bilirubine (BIL) values by maximum grade

End point title	Number of patients with abnormal Bilirubine (BIL) values by maximum grade[14]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. BIL – SCR G0; SE G3 = BIL with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[14] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
BIL - SCR G0; SE G0	24			
BIL - SCR G0; SE G1	0			
BIL - SCR G0; SE G2	0			
BIL - SCR G0; SE G3	0			
BIL - SCR G0; SE G4	0			
BIL - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Creatinine (CREA) values by maximum grade

End point title	Number of patients with abnormal Creatinine (CREA) values by maximum grade[15]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. CREA – SCR G0; SE G3 = CREA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[15] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
CREA - SCR G0; SE G0	20			
CREA - SCR G0; SE G1	2			
CREA - SCR G0; SE G2	0			
CREA - SCR G0; SE G3	0			
CREA - SCR G0; SE G4	0			
CREA - SCR G0; SE UNK	1			
CREA - SCR G1; SE G0	0			

CREA - SCR G1; SE G1	0			
CREA - SCR G1; SE G2	1			
CREA - SCR G1; SE G3	0			
CREA - SCR G1; SE UNK	0			
CREA - SCR G1; SE G4	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal gamma-glutamyl transpeptidase (GGT) values by maximum grade

End point title	Number of patients with abnormal gamma-glutamyl transpeptidase (GGT) values by maximum grade[16]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. GGT – SCR G0; SE G3 = GGT with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[16] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
GGT - SCR UNK; SE G0	0			
GGT - SCR UNK; SE G1	1			
GGT - SCR UNK; SE G2	0			
GGT - SCR UNK; SE G3	0			
GGT - SCR UNK; SE G4	0			
GGT - SCR UNK; SE UNK	0			
GGT - SCR G0; SE G0	17			
GGT - SCR G0; SE G1	2			
GGT - SCR G0; SE G2	1			
GGT - SCR G0; SE G3	0			
GGT - SCR G0; SE G4	0			
GGT - SCR G0; SE UNK	0			
GGT - SCR G1; SE G0	0			
GGT - SCR G1; SE G1	2			
GGT - SCR G1; SE G2	0			
GGT - SCR G1; SE G3	1			
GGT - SCR G1; SE G4	0			

GGT - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Hemoglobin (HGB) values by maximum grade

End point title	Number of patients with abnormal Hemoglobin (HGB) values by maximum grade[17]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. HGB – SCR G0; SE G3 = HGB with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[17] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
HGB - SCR G0; SE G0	14			
HGB - SCR G0; SE G1	9			
HGB - SCR G0; SE G2	0			
HGB - SCR G0; SE G3	0			
HGB - SCR G0; SE G4	0			
HGB - SCR G0; SE UNK	0			
HGB - SCR G1; SE G0	0			
HGB - SCR G1; SE G1	1			
HGB - SCR G1; SE G2	0			
HGB - SCR G1; SE G3	0			
HGB - SCR G1; SE G4	0			
HGB - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Hypercalcemia (HCA) values by maximum grade

End point title	Number of patients with abnormal Hypercalcemia (HCA) values by maximum grade[18]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. HCA – SCR G0; SE G3 = HCA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[18] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
HCA - SCR UNK; SE G0	3			
HCA - SCR UNK; SE G1	0			
HCA - SCR UNK; SE G2	0			
HCA - SCR UNK; SE G3	0			
HCA - SCR UNK; SE G4	0			
HCA - SCR UNK; SE UNK	0			
HCA - SCR G0; SE G0	15			
HCA - SCR G0; SE G1	3			
HCA - SCR G0; SE G2	0			
HCA - SCR G0; SE G3	0			
HCA - SCR G0; SE G4	0			
HCA - SCR G0; SE UNK	0			
HCA - SCR G1; SE G0	1			
HCA - SCR G1; SE G1	2			
HCA - SCR G1; SE G2	0			
HCA - SCR G1; SE G3	0			
HCA - SCR G1; SE G4	0			
HCA - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Hyperkalemia (HKA) values by maximum grade

End point title	Number of patients with abnormal Hyperkalemia (HKA) values by maximum grade[19]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. HKA – SCR G0; SE G3 = HKA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[19] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
HKA - SCR UNK; SE G0	2			
HKA - SCR UNK; SE G1	0			
HKA - SCR UNK; SE G2	0			
HKA - SCR UNK; SE G3	0			
HKA - SCR UNK; SE G4	0			
HKA - SCR UNK; SE UNK	0			
HKA - SCR G0; SE G0	17			
HKA - SCR G0; SE G1	3			
HKA - SCR G0; SE G2	1			
HKA - SCR G0; SE G3	0			
HKA - SCR G0; SE G4	0			
HKA - SCR G0; SE UNK	0			
HKA - SCR G1; SE G0	0			
HKA - SCR G1; SE G1	0			
HKA - SCR G1; SE G2	1			
HKA - SCR G1; SE G3	0			
HKA - SCR G1; SE G4	0			
HKA - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Hypernatremia (HNA) values by maximum grade

End point title	Number of patients with abnormal Hypernatremia (HNA) values by maximum grade[20]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus

baseline values, at Screening (SCR). [e.g. HNA – SCR G0; SE G3 = HNA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[20] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
HNA - SCR UNK; SE G0	2			
HNA - SCR UNK; SE G1	0			
HNA - SCR UNK; SE G2	0			
HNA - SCR UNK; SE G3	0			
HNA - SCR UNK; SE G4	0			
HNA - SCR UNK; SE UNK	0			
HNA - SCR G0; SE G0	19			
HNA - SCR G0; SE G1	3			
HNA - SCR G0; SE G2	0			
HNA - SCR G0; SE G3	0			
HNA - SCR G0; SE G4	0			
HNA - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal hypoalbuminemia(hAL) values by maximum grade

End point title	Number of patients with abnormal hypoalbuminemia(hAL) values by maximum grade[21]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. hAL – SCR G0; SE G3 = hAL with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[21] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
hAL - SCR UNK; SE G0	1			
hAL - SCR UNK; SE G1	1			
hAL - SCR UNK; SE G2	0			
hAL - SCR UNK; SE G3	0			
hAL - SCR UNK; SE G4	0			
hAL - SCR UNK; SE UNK	0			
hAL - SCR G0; SE G0	21			
hAL - SCR G0; SE G1	0			
hAL - SCR G0; SE G2	0			
hAL - SCR G0; SE G3	0			
hAL - SCR G0; SE G4	0			
hAL - SCR G0; SE UNK	1			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal hypocalcemia(hCA) values by maximum grade

End point title	Number of patients with abnormal hypocalcemia(hCA) values by maximum grade[22]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. hCA – SCR G0; SE G3 = hCA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[22] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
hCA - SCR UNK; SE G0	3			
hCA - SCR UNK; SE G1	0			
hCA - SCR UNK; SE G2	0			
hCA - SCR UNK; SE G3	0			
hCA - SCR UNK; SE G4	0			

hCA - SCR UNK; SE UNK	0			
hCA - SCR G0; SE G0	18			
hCA - SCR G0; SE G1	2			
hCA - SCR G0; SE G2	0			
hCA - SCR G0; SE G3	0			
hCA - SCR G0; SE G4	0			
hCA - SCR G0; SE UNK	0			
hCA - SCR G1; SE G0	0			
hCA - SCR G1; SE G1	1			
hCA - SCR G1; SE G2	0			
hCA - SCR G1; SE G3	0			
hCA - SCR G1; SE G4	0			
hCA - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal hypokalemia (hKA) values by maximum grade

End point title	Number of patients with abnormal hypokalemia (hKA) values by maximum grade[23]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. hKA – SCR G0; SE G3 = hKA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[23] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
hKA - SCR UNK; SE G0	2			
hKA - SCR UNK; SE G1	0			
hKA - SCR UNK; SE G2	0			
hKA - SCR UNK; SE G3	0			
hKA - SCR UNK; SE G4	0			
hKA - SCR UNK; SE UNK	0			
hKA - SCR G0; SE G0	19			
hKA - SCR G0; SE G1	3			
hKA - SCR G0; SE G2	0			

hKA - SCR G0; SE G3	0			
hKA - SCR G0; SE G4	0			
hKA - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal hyponatremia (hNA) values by maximum grade

End point title	Number of patients with abnormal hyponatremia (hNA) values by maximum grade[24]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. hNA – SCR G0; SE G3 = hNA with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[24] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
hNA - SCR UNK; SE G0	2			
hNA - SCR UNK; SE G1	0			
hNA - SCR UNK; SE G2	0			
hNA - SCR UNK; SE G3	0			
hNA - SCR UNK; SE G4	0			
hNA - SCR UNK; SE UNK	0			
hNA - SCR G0; SE G0	18			
hNA - SCR G0; SE G1	4			
hNA - SCR G0; SE G2	0			
hNA - SCR G0; SE G3	0			
hNA - SCR G0; SE G4	0			
hNA - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Leukocytes (LEU) values by maximum grade

End point title	Number of patients with abnormal Leukocytes (LEU) values by maximum grade[25]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. LEU – SCR G0; SE G3 = LEU with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[25] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
LEU - SCR G0; SE G0	21			
LEU - SCR G0; SE G1	2			
LEU - SCR G0; SE G2	0			
LEU - SCR G0; SE G3	0			
LEU - SCR G0; SE G4	0			
LEU - SCR G0; SE UNK	0			
LEU - SCR G1; SE G0	1			
LEU - SCR G1; SE G1	0			
LEU - SCR G1; SE G2	0			
LEU - SCR G1; SE G3	0			
LEU - SCR G1; SE G4	0			
LEU - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Lymphopenia (LYM) values by maximum grade

End point title	Number of patients with abnormal Lymphopenia (LYM) values by maximum grade[26]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. LYM – SCR G0; SE G3 = LYM with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[26] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
LYM - SCR G0; SE G0	17			
LYM - SCR G0; SE G1	0			
LYM - SCR G0; SE G2	3			
LYM - SCR G0; SE G3	0			
LYM - SCR G0; SE G4	0			
LYM - SCR G0; SE UNK	0			
LYM - SCR G1; SE G0	1			
LYM - SCR G1; SE G1	3			
LYM - SCR G1; SE G2	0			
LYM - SCR G1; SE G3	0			
LYM - SCR G1; SE G4	0			
LYM - SCR G1; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Neutrophils (NEU) values by maximum grade

End point title	Number of patients with abnormal Neutrophils (NEU) values by maximum grade[27]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. NEU – SCR G0; SE G3 = NEU with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[27] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
NEU - SCR G0; SE G0	21			
NEU - SCR G0; SE G1	2			
NEU - SCR G0; SE G2	1			
NEU - SCR G0; SE G3	0			
NEU - SCR G0; SE G4	0			
NEU - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with abnormal Platelets(PLT) values by maximum grade

End point title	Number of patients with abnormal Platelets(PLT) values by maximum grade[28]

End point description:

The status of each patient was collected and graded according to the Common Terminology Criteria Adverse Event (CTCAE v.03) terminology. Gradings were: G0 (normal value), G1 (Mild abnormality), G2 (Moderate abnormality), G3 (Severe abnormality), G4 (Life-threatening/Disabling abnormality), Unknown abnormality (UNK). The post-treatment values, at Study End (SE) were presented versus baseline values, at Screening (SCR). [e.g. PLT – SCR G0; SE G3 = PLT with no abnormality/normal value at screening and with Severe abnormality at study end].

End point type	Primary

End point timeframe:

From Screening and up to Week 11 (Cycle 1), Week 30 (Cycle 2), Week 52 (Cycle 3), and Study End at Year 4 (Cycle 4)

Notes:

[28] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
PLT - SCR G0; SE G0	24			
PLT - SCR G0; SE G1	0			
PLT - SCR G0; SE G2	0			
PLT - SCR G0; SE G3	0			
PLT - SCR G0; SE G4	0			
PLT - SCR G0; SE UNK	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with any AE(s) and with AEs by maximum grade, related to treatment administration

End point title	Number of patients with any AE(s) and with AEs by maximum grade, related to treatment administration[29]

End point description:

An AE was any untoward medical occurrence in a patient or clinical investigation subject, temporally associated with the use of a medicinal product, whether or not considered related to the medicinal product. AEs reported are here below tabulated irrespective of grade (any), as well as graded by maximum grade reported according to the Common Terminology Criteria (CTC) Adverse event terminology, version 3.0. Maximum grade reported and tabulated were Grade 1 (G1) – Mild AE, G2 – Moderate AE, G3 – Severe AE, G4 – Life threatening/Disabling AE and G5 – Death related to AE.

End point type	Primary

End point timeframe:

Within the 31-day follow-up period post treatment administration.

Notes:

[29] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Patients with any AEs	23			
Patients with G1 AEs	11			
Patients with G2 AEs	12			
Patients with G3 AEs	0			
Patients with G4 AEs	0			
Patients with G5 AEs	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with any adverse events (AEs) and with AEs by maximum grade

End point title	Number of patients with any adverse events (AEs) and with AEs by maximum grade[30]

End point description:

An AE was any untoward medical occurrence in a patient or clinical investigation subject, temporally associated with the use of a medicinal product, whether or not considered related to the medicinal product. AEs reported are here below tabulated irrespective of grade (any), as well as graded by maximum grade reported according to the CTC Adverse event terminology, version 3.0. Maximum grade reported and tabulated were Grade 1 (G1) – Mild AE, G2 – Moderate AE, G3 – Severe AE, G4 – Life threatening/Disabling AE and G5 – Death related to AE.

End point type	Primary

End point timeframe:

Within the 31-day follow-up period post treatment administration.

Notes:

[30] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Patients with any AEs	24			
Patients with G1 AEs	6			
Patients with G2 AEs	15			
Patients with G3 AEs	3			
Patients with G4 AEs	0			
Patients with G5 AEs	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with any serious adverse events (SAEs) and with SAEs by maximum grade

End point title	Number of patients with any serious adverse events (SAEs) and with SAEs by maximum grade[31]

End point description:

SAEs include medical occurrences that result in death, are life threatening, require hospitalization or prolongation of hospitalization or result in disability/incapacity, is a congenital anomaly/birth defect in the offspring of a patient. Events which were part of the natural course of the disease under study were captured as part of the clinical activity outcome variables in this study; therefore did not need to be reported as SAEs. Progression/recurrence of the tumor was recorded as part of the clinical assessment data collection, and deaths due to progressive disease was recorded on a specific form, but not as an SAE. SAEs reported are here below tabulated irrespective of grade (any), as well as graded by maximum grade reported according to the CTC Adverse event terminology, version 3.0. Maximum grade reported and tabulated were Grade 1 (G1) – Mild SAE, G2 – Moderate SAE, G3 – Severe SAE, G4 – Life threatening/Disabling SAE and G5 – Death related to SAE.

End point type	Primary

End point timeframe:

Within the 31-day follow-up period post treatment administration.

Notes:

[31] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Patients with any SAEs	2			
Patients with G1 SAEs	0			
Patients with G2 SAEs	1			
Patients with G3 SAEs	1			
Patients with G4 SAEs	0			

Patients with G5 SAEs	0			

Statistical analyses

No statistical analyses for this end point

Primary: Number of patients with any serious adverse events (SAEs) and with SAEs by maximum grade, related to treatment administration

End point title	Number of patients with any serious adverse events (SAEs) and with SAEs by maximum grade, related to treatment administration[32]

End point description:

SAEs include medical occurrences that result in death, are life threatening, require hospitalization or prolongation of hospitalization or result in disability/incapacity, is a congenital anomaly/birth defect in the offspring of a patient. Events which were part of the natural course of the disease under study were captured as part of the clinical activity outcome variables in this study; therefore did not need to be reported as SAEs. Progression/recurrence of the tumor was recorded as part of the clinical assessment data collection, and deaths due to progressive disease was recorded on a specific form, but not as an SAE. SAEs reported are here below tabulated irrespective of grade (any), as well as graded by maximum grade reported according to the CTC Adverse event terminology, version 3.0. Maximum grade reported and tabulated were Grade 1 (G1) – Mild SAE, G2 – Moderate SAE, G3 – Severe SAE, G4 – Life threatening/Disabling SAE and G5 – Death related to SAE.

End point type	Primary

End point timeframe:

Within the 31-day follow-up period post treatment administration.

Notes:

[32] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	24			
Units: Patients				
Patients with any SAEs	0			
Patients with G1 SAEs	0			
Patients with G2 SAEs	0			
Patients with G3 SAEs	0			
Patients with G4 SAEs	0			
Patients with G5 SAEs	0			

Statistical analyses

No statistical analyses for this end point

Primary: Geometric Mean titers of Anti-MAGE-A3 specific CD4+ and CD8+ T-cells concentrations after immunization

End point title	Geometric Mean titers of Anti-MAGE-A3 specific CD4+ and

	CD8+ T-cells concentrations after immunization[33]

End point description:

This endpoint presents the geometric mean concentration, expressed in titers, of anti-MAGE-A3 specific CD4+ and CD8+ T-cells. These specific T-cells included the cluster of differentiation 4+ (CD4+) and CD8+ T-cells producing cytokines Tumor Necrosis Factor-alpha (TNF-α) and/or Interferon-gamma (INF-γ).

End point type	Primary

End point timeframe:

From Pre-treatment (up to 4 weeks before first treatment) to Concluding visit (CCL: at Week 196 + 30 to 37 days for patients completing the treatment, 1 month after the last Dose administered for patients withdrawn from study treatment before completion)

Notes:

[33] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: This outcome was descriptive, hence no statistical analyses were performed.

End point values	MAGE-A3 Group			
Subject group type	Reporting group			
Number of subjects analysed	18			
Units: Titers				
geometric mean (confidence interval 95%)				
CD4.TNFα (+) + IFNγ (+) PRE (N=17)	1.05 (0.98 to 1.11)			
CD8.TNFα (+) + IFNγ (+) PRE (N=18)	1.01 (0.99 to 1.03)			
CD4.TNFα (+) + IFNγ (+) (W5) (N=17)	3.36 (2.2 to 5.13)			
CD8.TNFα (+) + IFNγ (+) (W5) (N=17)	1 (1 to 1)			
CD4.TNFα (+) + IFNγ (+) (W13) (N=14)	5.35 (2.42 to 11.79)			
CD8.TNFα (+) + IFNγ (+) (W13) (N=15)	1 (1 to 1)			
CD4.TNFα (+) + IFNγ (+) (W32) (N=10)	5.29 (2.09 to 13.37)			
CD8.TNFα (+) + IFNγ (+) (W32) (N=10)	1.04 (0.99 to 1.09)			
CD4.TNFα (+) + IFNγ (+) (W54) (N=5)	2.83 (1.04 to 7.68)			
CD8.TNFα (+) + IFNγ (+) (W54) (N=5)	1 (1 to 1)			
CD4.TNFα (+) + IFNγ (+) (W78) (N=2)	2.62 (0.07 to 92.58)			
CD8.TNFα (+) + IFNγ (+) (W78) (N=2)	1 (1 to 1)			
CD4.TNFα (+) + IFNγ (+) (W102) (N=3)	2.13 (0.29 to 15.91)			
CD8.TNFα (+) + IFNγ (+) (W102) (N=3)	1.05 (0.86 to 1.28)			
CD4.TNFα (+) + IFNγ (+) CCL (N=2)	2.42 (0 to 185163.6)			
CD8.TNFα (+) + IFNγ (+) CCL (N=2)	1 (1 to 1)			

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information	
Timeframe for reporting adverse events: Unsolicited AEs during the 31-day post-vaccination (Days 0-30), SAEs during the entire study period (Day 0 - Year 4).	
Assessment type	Non-systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	18.0

Reporting groups	
Reporting group title	MAGE-A3 Group
Reporting group description: Patients planned to receive intramuscularly up to 24 doses of MAGE-A3 ASCI (the study product), in 4 cycles.	

Serious adverse events	MAGE-A3 Group		
Total subjects affected by serious adverse events			
subjects affected / exposed	2 / 24 (8.33%)		
number of deaths (all causes)	0		
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Metastatic malignant melanoma			
subjects affected / exposed	1 / 24 (4.17%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cardiac disorders			
Cardiac disorder			
subjects affected / exposed	1 / 24 (4.17%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		

Frequency threshold for reporting non-serious adverse events: 5 %

Non-serious adverse events	MAGE-A3 Group		
Total subjects affected by non-serious adverse events			
subjects affected / exposed	23 / 24 (95.83%)		
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			

Cancer pain			
subjects affected / exposed	2 / 24 (8.33%)		
occurrences (all)	2		
Nervous system disorders			
Headache			
subjects affected / exposed	7 / 24 (29.17%)		
occurrences (all)	31		
General disorders and administration site conditions			
Injection site pain			
subjects affected / exposed	12 / 24 (50.00%)		
occurrences (all)	40		
Injection site reaction			
subjects affected / exposed	12 / 24 (50.00%)		
occurrences (all)	50		
Pyrexia			
subjects affected / exposed	10 / 24 (41.67%)		
occurrences (all)	31		
Asthenia			
subjects affected / exposed	9 / 24 (37.50%)		
occurrences (all)	19		
Fatigue			
subjects affected / exposed	7 / 24 (29.17%)		
occurrences (all)	13		
Influenza like illness			
subjects affected / exposed	7 / 24 (29.17%)		
occurrences (all)	40		
Chills			
subjects affected / exposed	4 / 24 (16.67%)		
occurrences (all)	8		
Injection site erythema			
subjects affected / exposed	4 / 24 (16.67%)		
occurrences (all)	7		
Discomfort			
subjects affected / exposed	3 / 24 (12.50%)		
occurrences (all)	10		
Administration site pain			

subjects affected / exposed occurrences (all) Injection site induration subjects affected / exposed occurrences (all) Ulcer subjects affected / exposed occurrences (all)	2 / 24 (8.33%) 4 2 / 24 (8.33%) 7 2 / 24 (8.33%) 2		
Gastrointestinal disorders Nausea subjects affected / exposed occurrences (all) Diarrhoea subjects affected / exposed occurrences (all) Vomiting subjects affected / exposed occurrences (all)	 7 / 24 (29.17%) 17 2 / 24 (8.33%) 2 2 / 24 (8.33%) 2		
Respiratory, thoracic and mediastinal disorders Cough subjects affected / exposed occurrences (all) Dyspnoea subjects affected / exposed occurrences (all)	 3 / 24 (12.50%) 3 2 / 24 (8.33%) 2		
Skin and subcutaneous tissue disorders Erythema subjects affected / exposed occurrences (all) Hyperhidrosis subjects affected / exposed occurrences (all) Pruritus subjects affected / exposed occurrences (all)	 2 / 24 (8.33%) 2 2 / 24 (8.33%) 2 2 / 24 (8.33%) 7		
Musculoskeletal and connective tissue disorders			

Arthralgia subjects affected / exposed occurrences (all) Myalgia subjects affected / exposed occurrences (all) Back pain subjects affected / exposed occurrences (all) Groin pain subjects affected / exposed occurrences (all)	3 / 24 (12.50%) 3 3 / 24 (12.50%) 18 2 / 24 (8.33%) 2 2 / 24 (8.33%) 2		
Infections and infestations Skin infection subjects affected / exposed occurrences (all) Upper respiratory tract infection subjects affected / exposed occurrences (all)	2 / 24 (8.33%) 2 2 / 24 (8.33%) 2		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
02 March 2009	The changes following this amendment concern: • The possibility of taking a new tumor biopsy in case the results of the analysis of the biopsies originally taken are inconclusive. • The description of the AJCC staging system for cutaneous melanoma • The AJCC staging system for cutaneous melanoma • The time window for tumor imaging at screening • The expected time for completing patient recruitment

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported