Clinical trial results:
A Phase III, Randomized, Double-Blind, Placebo-Controlled, Adaptive Design Study of the Efficacy, Safety, and Tolerability of a Single Infusion of MK-3415 (Human Monoclonal Antibody to Clostridium difficile toxin A), MK-6072 (Human Monoclonal Antibody to Clostridium difficile toxin B), and MK-3415A (Human Monoclonal Antibodies to Clostridium difficile toxin A and toxin B) in Patients Receiving Antibiotic Therapy for Clostridium difficile Infection (MODIFY I)

Due to the EudraCT – Results system being out of service between 31 July 2015 and 12 January 2016, these results have been published in compliance with revised timelines.

Summary

EudraCT number	2011-004590-90
Trial protocol	DE ES CZ BE DK PT AT IT GB
Global end of trial date	09 December 2014

Results information

Result version number	v1 (current)
This version publication date	06 February 2016
First version publication date	06 February 2016

Trial information

Trial identification

Sponsor protocol code	3415A-001

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT01241552
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Merck Sharp & Dohme Corp.
Sponsor organisation address	2000 Galloping Hill Road, Kenilworth, NJ, United States, 07033
Public contact	Clinical Trials Disclosure, Merck Sharp & Dohme Corp., ClinicalTrialsDisclosure@merck.com
Scientific contact	Clinical Trials Disclosure, Merck Sharp & Dohme Corp., ClinicalTrialsDisclosure@merck.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	09 December 2014
Is this the analysis of the primary completion data?	Yes
Primary completion date	09 December 2014
Global end of trial reached?	Yes
Global end of trial date	09 December 2014
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

Primary Objective #1 (at interim analysis) and #2 (at final analysis): To determine if treatment with a single infusion of combined monoclonal antibody therapy (MK-3415A) with standard of care therapy decreases the proportion of participants with CDI recurrence over a period of 12 weeks as compared to treatment with a single infusion of individual monoclonal antibody therapy (MK-3415 or MK-6072) with standard of care therapy.

Primary Objective #3: To determine if treatment with a single infusion of monoclonal antibody therapy with standard of care therapy (combined monoclonal antibody therapy [MK-3415A] and possibly the separate individual monoclonal antibody therapy [MK-3415 and/or MK-6072]) decreases the proportion of participants with CDI recurrence over a period of 12 weeks as compared to treatment with a single infusion of placebo with standard of care therapy.

Protection of trial subjects:

This study was conducted in conformance with Good Clinical Practice standards and applicable country and/or local statutes and regulations regarding ethical committee review, informed consent, and the protection of human subjects participating in biomedical research.

Background therapy:

Oral SOC antibiotic therapy (metronidazole, vancomycin, or fidaxomicin) for a primary or recurrent episode of CDI.

Evidence for comparator: -

Actual start date of recruitment	10 October 2011
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	Australia: 52
Country: Number of subjects enrolled	Austria: 18
Country: Number of subjects enrolled	Belgium: 28
Country: Number of subjects enrolled	Brazil: 11
Country: Number of subjects enrolled	Canada: 89
Country: Number of subjects enrolled	Chile: 55
Country: Number of subjects enrolled	Colombia: 23
Country: Number of subjects enrolled	Czech Republic: 34
Country: Number of subjects enrolled	Denmark: 69

Country: Number of subjects enrolled	Germany: 31
Country: Number of subjects enrolled	Israel: 85
Country: Number of subjects enrolled	Italy: 82
Country: Number of subjects enrolled	Mexico: 8
Country: Number of subjects enrolled	New Zealand: 22
Country: Number of subjects enrolled	Portugal: 25
Country: Number of subjects enrolled	South Africa: 10
Country: Number of subjects enrolled	Spain: 85
Country: Number of subjects enrolled	United Kingdom: 37
Country: Number of subjects enrolled	United States: 688
Worldwide total number of subjects	1452
EEA total number of subjects	409

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)	721
From 65 to 84 years	581
85 years and over	150

Subject disposition

Recruitment

Recruitment details: -

Pre-assignment

Screening details:

Participants 18 years of age or older, with a diagnosis of CDI were enrolled in this trial.

Period 1

Period 1 title	Treatment Period 1 (overall period)
Is this the baseline period?	Yes
Allocation method	Randomised - controlled
Blinding used	Double blind
Roles blinded	Subject, Investigator, Carer, Assessor

Arms

Are arms mutually exclusive?	Yes
Arm title	MK-3415 + SOC

Arm description:

Single intravenous (IV) infusion of 10 mg/kg MK-3415 + SOC for CDI

Arm type	Experimental
Investigational medicinal product name	MK-3415
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Infusion
Routes of administration	Intravenous use

Dosage and administration details:

A single IV infusion of MK-3415 (10 mg/kg of monoclonal antibody to Clostridium difficile Toxin A)

Investigational medicinal product name	SOC
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

SOC for CDI was antibiotic therapy consisting of metronidazole, vancomycin, or fidaxomicin prescribed for 10 to 14 days beginning prior to or on the day of study drug

Arm title	MK-6072+ SOC

Arm description:

Single IV infusion of 10 mg/kg MK-6072 + Standard of Care for CDI

Arm type	Experimental
Investigational medicinal product name	MK-6072
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Infusion
Routes of administration	Intravenous use

Dosage and administration details:

A single IV infusion of MK-6072 (10 mg/kg of monoclonal antibody to Clostridium difficile Toxin B)

Investigational medicinal product name	SOC
Investigational medicinal product code	
Other name	

Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

SOC for CDI was antibiotic therapy consisting of metronidazole, vancomycin, or fidaxomicin prescribed for 10 to 14 days beginning prior to or on the day of study drug

Arm title	MK-3415A + SOC

Arm description:

Single IV infusion of 10 mg/kg MK-3415A + Standard of Care for CDI

Arm type	Experimental
Investigational medicinal product name	SOC
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

SOC for CDI was antibiotic therapy consisting of metronidazole, vancomycin, or fidaxomicin prescribed for 10 to 14 days beginning prior to or on the day of study drug

Investigational medicinal product name	MK-3415A
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Infusion
Routes of administration	Intravenous use

Dosage and administration details:

A single IV infusion of MK3415A (10 mg/kg of monoclonal antibody to Clostridium difficile Toxin A and 10mg/kg of monoclonal antibody to Clostridium difficile Toxin B)

Arm title	Placebo Comparator + SOC

Arm description:

Normal saline infusion (0.9% sodium chloride) + Standard of Care for CDI

Arm type	Placebo
Investigational medicinal product name	Placebo
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Infusion
Routes of administration	Intravenous use

Dosage and administration details:

A single IV infusion of normal saline (0.9% sodium chloride)

Investigational medicinal product name	SOC
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

SOC for CDI was antibiotic therapy consisting of metronidazole, vancomycin, or fidaxomicin prescribed for 10 to 14 days beginning prior to or on the day of study drug

Number of subjects in period 1	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC
Started	242	403	403
Treated	235	392	388
Completed	201	340	343
Not completed	41	63	60
Consent withdrawn by subject	7	15	14
Physician decision	2	4	2
Adverse event, non-fatal	1	1	-
Death	26	30	20
Technical Problems	2	-	2
Progressive Disease	-	-	1
Lost to follow-up	2	11	15
Lack of efficacy	1	-	-
Protocol deviation	-	2	6

Number of subjects in period 1	Placebo Comparator + SOC
Started	404
Treated	397
Completed	340
Not completed	64
Consent withdrawn by subject	15
Physician decision	3
Adverse event, non-fatal	-
Death	25
Technical Problems	2
Progressive Disease	2
Lost to follow-up	16
Lack of efficacy	-
Protocol deviation	1

Baseline characteristics

Reporting groups	
Reporting group title	MK-3415 + SOC
Reporting group description:	
Single intravenous (IV) infusion of 10 mg/kg MK-3415 + SOC for CDI	
Reporting group title	MK-6072+ SOC
Reporting group description:	
Single IV infusion of 10 mg/kg MK-6072 + Standard of Care for CDI	
Reporting group title	MK-3415A + SOC
Reporting group description:	
Single IV infusion of 10 mg/kg MK-3415A + Standard of Care for CDI	
Reporting group title	Placebo Comparator + SOC
Reporting group description:	
Normal saline infusion (0.9% sodium chloride) + Standard of Care for CDI	

Reporting group values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC
Number of subjects	242	403	403
Age Categorical			
Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	117	210	195
From 65-84 years	92	153	178
85 years and over	33	40	30
Age Continuous			
Units: years			
arithmetic mean	64.2	61.1	62.5
standard deviation	± 16.8	± 18.5	± 17.8
Gender Categorical			
Units: Subjects			
Female	137	238	224
Male	105	165	179

Reporting group values	Placebo Comparator + SOC	Total	
Number of subjects	404	1452	
Age Categorical			
Units: Subjects			
In utero	0	0	
Preterm newborn infants (gestational age < 37 wks)	0	0	
Newborns (0-27 days)	0	0	

Infants and toddlers (28 days-23 months)	0	0	
Children (2-11 years)	0	0	
Adolescents (12-17 years)	0	0	
Adults (18-64 years)	199	721	
From 65-84 years	158	581	
85 years and over	47	150	
Age Continuous			
Units: years			
arithmetic mean	62.9		
standard deviation	± 18.3	-	
Gender Categorical			
Units: Subjects			
Female	230	829	
Male	174	623	

End points

End points reporting groups	
Reporting group title	MK-3415 + SOC
Reporting group description:	
Single intravenous (IV) infusion of 10 mg/kg MK-3415 + SOC for CDI	
Reporting group title	MK-6072+ SOC
Reporting group description:	
Single IV infusion of 10 mg/kg MK-6072 + Standard of Care for CDI	
Reporting group title	MK-3415A + SOC
Reporting group description:	
Single IV infusion of 10 mg/kg MK-3415A + Standard of Care for CDI	
Reporting group title	Placebo Comparator + SOC
Reporting group description:	
Normal saline infusion (0.9% sodium chloride) + Standard of Care for CDI	

Primary: Percentage of participants with CDI recurrence	
End point title	Percentage of participants with CDI recurrence
End point description:	
CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.	
End point type	Primary
End point timeframe:	
Up to 12 weeks	

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	232	386	383	395
Units: Percentage of participants				
number (not applicable)	25.9	17.4	15.9	27.6

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	627
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.3182 [1]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-1.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-8.6
upper limit	5.5

Notes:

[1] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	781
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0003 [2]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-10.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-15.9
upper limit	-4.3

Notes:

[2] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	778
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001 [3]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-11.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-17.4
upper limit	-5.9

Notes:

[3] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - MK-3415 + SOC	
Comparison groups	MK-3415 + SOC v MK-3415A + SOC
Number of subjects included in analysis	615
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0013 [4]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-9.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.9
upper limit	-3.4

Notes:

[4] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - MK-6072 + SOC	
Comparison groups	MK-6072+ SOC v MK-3415A + SOC
Number of subjects included in analysis	769
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.2997 [5]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-1.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-6.7
upper limit	3.9

Notes:

[5] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Primary: Percentage of participants with one or more Adverse Events (AEs) during 4 weeks following infusion

End point title	Percentage of participants with one or more Adverse Events (AEs) during 4 weeks following infusion

End point description:

An AE is defined as any untoward medical occurrence in a clinical investigation participant administered a pharmaceutical product and which does not necessarily have to have a causal relationship with this treatment. An AE can therefore be any unfavorable and unintended signs (including an abnormal laboratory finding, for example), symptom, or disease temporally associated with the use of a medicinal product or protocol-specific procedure, whether or not considered related to the medicinal product or protocol-specific procedure. Any worsening (i.e., any clinically significant adverse change in frequency

and/or intensity) of a preexisting condition that is temporally associated with the use of the Sponsor's product, is also an AE. The population analyzed consisted of all randomized participants who received infusion of study medication.

End point type	Primary
End point timeframe:	
Up to 28 days	

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	67.2	65.4	59.7	62

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.185
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	5.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-2.5
upper limit	12.8

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.323
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	3.4

Confidence interval	
level	95 %
sides	2-sided
lower limit	-3.3
upper limit	10.1

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.507
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	-2.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-9.1
upper limit	4.5

Primary: Percentage of participants with any drug-related AE during 4 weeks following infusion

End point title	Percentage of participants with any drug-related AE during 4 weeks following infusion

End point description:

An AE is defined as any untoward medical occurrence in a clinical investigation participant administered a pharmaceutical product and which does not necessarily have to have a causal relationship with this treatment. An AE can therefore be any unfavorable and unintended signs (including an abnormal laboratory finding, for example), symptom, or disease temporally associated with the use of a medicinal product or protocol-specific procedure, whether or not considered related to the medicinal product or protocol-specific procedure. Any worsening (i.e., any clinically significant adverse change in frequency and/or intensity) of a preexisting condition that is temporally associated with the use of the Sponsor's product, is also an AE. A drug-related AE was an AE determined by the investigator to be related to the drug. The population analyzed consisted of all randomized participants who received infusion of study medication.

End point type	Primary

End point timeframe:

Up to 28 days

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	7.2	8.2	6.2	5

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.246
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	2.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-1.5
upper limit	6.7

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.069
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	3.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.3
upper limit	6.8

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.464
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	1.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-2.1
upper limit	4.6

Primary: Percentage of participants with any serious adverse events (SAEs) during 4 weeks following infusion

End point title	Percentage of participants with any serious adverse events (SAEs) during 4 weeks following infusion

End point description:

A SAE is any AE occurring at any dose or during any use of Sponsor's product that: results in death; or is life threatening; or results in a persistent or significant disability/incapacity; or results in or prolongs an existing inpatient hospitalization; or is a congenital anomaly/birth defect; or is a cancer, or is associated with an overdose (whether accidental or intentional); or is other important medical events. The population analyzed consisted of all randomized participants who received infusion of study medication

End point type	Primary

End point timeframe:

Up to 28 days

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	27.7	21.5	14.7	20

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC

Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.027
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	7.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.9
upper limit	14.7

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.594
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	1.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-4.1
upper limit	7.2

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.051
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	-5.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-10.6
upper limit	0

Primary: Percentage of participants with any serious drug-related AEs (SAEs) during 4 weeks following infusion

End point title	Percentage of participants with any serious drug-related AEs (SAEs) during 4 weeks following infusion

End point description:

A SAE is any AE occurring at any dose or during any use of Sponsor's product that: results in death; or is life threatening; or results in a persistent or significant disability/incapacity; or results in or prolongs an existing inpatient hospitalization; or is a congenital anomaly/birth defect; or is a cancer, or is associated with an overdose (whether accidental or intentional); or is other important medical events. A serious drug-related AE was an SAE determined by the investigator to be related to the drug. The population analyzed consisted of all randomized participants who received infusion of study medication

End point type	Primary

End point timeframe:

Up to 28 days

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	1.3	1	0.5	0.3

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC

Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.115
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.3
upper limit	3.5

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.17
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	0.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.5
upper limit	2.4

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.544
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	0.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.9
upper limit	1.6

Primary: Percentage of participants who discontinued study medication due to an AE during 4 weeks following infusion

End point title	Percentage of participants who discontinued study medication due to an AE during 4 weeks following infusion

End point description:

An AE is defined as any untoward medical occurrence in a clinical investigation participant administered a pharmaceutical product and which does not necessarily have to have a causal relationship with this treatment. An AE can therefore be any unfavorable and unintended signs (including an abnormal laboratory finding, for example), symptom, or disease temporally associated with the use of a medicinal product or protocol-specific procedure, whether or not considered related to the medicinal product or protocol-specific procedure. Any worsening (i.e., any clinically significant adverse change in frequency and/or intensity) of a preexisting condition that is temporally associated with the use of the Sponsor’s product, is also an AE. The population analyzed consisted of all randomized participants who received infusion of study medication.

End point type	Primary

End point timeframe:
Up to 28 days

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	0.4	0.3	0	0

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.192
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	0.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.5
upper limit	2.4

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.311
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	0.3

Confidence interval	
level	95 %
sides	2-sided
lower limit	-0.7
upper limit	1.4

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
P-value	> 0.999
Method	Miettinen and Nurminen
Parameter estimate	Percentage Difference
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	-1
upper limit	1

Secondary: Percentage of participants with infusion-specific AEs

End point title	Percentage of participants with infusion-specific AEs

End point description:

Infusion-specific AEs included local infusion site AEs; and systemic AEs which include nausea, vomiting, chills, fatigue, feeling hot, infusion site conditions (bruising, coldness, erythema, extravasation, pain, phlebitis, pruritus), pyrexia, arthralgia, musculoskeletal pain, myalgia, dizziness, headache, dysphonia, nasal congestion, pruritus, rash, pruritic rash, urticaria, flushing, hot flush, hypertension, andhypotension. The population analyzed consisted of all randomized participants who received infusion of study medication.

End point type	Secondary

End point timeframe:

Up to 24 hours

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	235	390	387	400
Units: Percentage of participants				
number (not applicable)	11.1	11.8	8.8	7.5

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	635
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	3.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-1
upper limit	8.7

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	790
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	4.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.2
upper limit	8.5

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	787
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	1.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-2.6
upper limit	5.2

Secondary: Percentage of participants with Global Cure

End point title	Percentage of participants with Global Cure

End point description:

Global Cure is defined as the clinical cure of the initial CDI episode and no CDI recurrence through Week 12. Clinical cure is defined as participants who received ≤ 14 day regimen of SOC therapy and have no diarrhea (≤2 loose stools per 24 hours) for two consecutive days following completion of SOC therapy for the baseline CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	232	386	383	395
Units: Percentage of participants				
number (not applicable)	47	60.1	58.7	55.2

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Adjusted Difference: MK-3415 + SOC - Placebo Comparator + SOC

Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	627
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.9775 [6]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-8.3

Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.3
upper limit	-0.2

Notes:

[6] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	781
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0861 [7]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	4.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-2.1
upper limit	11.7

Notes:

[7] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	778
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.1646 [8]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	3.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-3.5
upper limit	10.4

Notes:

[8] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - MK-3415 + SOC	
Comparison groups	MK-3415 + SOC v MK-3415A + SOC

Number of subjects included in analysis	615
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0025 [9]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	11.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	3.5
upper limit	19.7

Notes:

[9] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Adjusted Difference: MK-3415A + SOC - MK-6072 + SOC

Comparison groups	MK-6072+ SOC v MK-3415A + SOC
Number of subjects included in analysis	769
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.6532 [10]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-1.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-8.3
upper limit	5.5

Notes:

[10] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Secondary: Percentage of participants with CDI recurrence in those with clinical cure of the initial CDI episode

End point title	Percentage of participants with CDI recurrence in those with clinical cure of the initial CDI episode

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. Clinical cure is defined as participants who received ≤ 14 day regimen of SOC therapy and have no diarrhea (≤2 loose stools per 24 hours) for two consecutive days following completion of SOC therapy for the baseline CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	169	299	286	327
Units: Percentage of participants				
number (not applicable)	35.5	22.4	21.3	33.3

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	496
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.6505 [11]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	1.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-6.9
upper limit	10.7

Notes:

[11] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	626
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0013 [12]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-10.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-17.7
upper limit	-3.8

Notes:

[12] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	613
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0006 [13]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-11.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-18.6
upper limit	-4.7

Notes:

[13] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - MK-3415 + SOC	
Comparison groups	MK-3415 + SOC v MK-3415A + SOC
Number of subjects included in analysis	455
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0007 [14]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-13.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-22.5
upper limit	-5.2

Notes:

[14] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Adjusted Difference: MK-3415A + SOC - MK-6072 + SOC	
Comparison groups	MK-6072+ SOC v MK-3415A + SOC

Number of subjects included in analysis	585
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.3906 [15]
Method	Miettinen and Nurminen
Parameter estimate	Adjusted Difference
Point estimate	-1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-7.7
upper limit	5.8

Notes:

[15] - One sided p-value based on the Miettinen and Nurminen method stratified by SOC therapy (metronidazole vs. vancomycin vs. fidaxomicin) and hospitalization status (inpatient vs. outpatient).

Secondary: Percentage of participants ≥ 65 years of age at study entry with CDI recurrence

End point title	Percentage of participants ≥ 65 years of age at study entry with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	122	185	200	199
Units: Percentage of participants				
number (not applicable)	26.2	15.1	17	33.2

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC

Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	321
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-6.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.8
upper limit	3.5

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	384
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-18
Confidence interval	
level	95 %
sides	2-sided
lower limit	-26.3
upper limit	-9.6

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	399
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-16.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-24.5
upper limit	-7.7

Secondary: Percentage of participants with a history of CDI in the 6 months prior to enrollment with CDI recurrence

End point title	Percentage of participants with a history of CDI in the 6

	months prior to enrollment with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	69	103	96	109
Units: Percentage of participants				
number (not applicable)	33.3	26.2	25	39.4

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	178
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-6.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-20.1
upper limit	8.6

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC

Number of subjects included in analysis	212
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-13.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-25.5
upper limit	-0.5

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	205
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-14.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-26.8
upper limit	-1.6

Secondary: Percentage of participants with clinically severe CDI at study entry with CDI recurrence

End point title	Percentage of participants with clinically severe CDI at study entry with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. With clinically severe CDI is defined as a Zar Score ≥ 2. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	31	67	62	60
Units: Percentage of participants				
number (not applicable)	25.8	10.4	12.9	25

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	91
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	0.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.9
upper limit	21

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	127
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-14.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-28.3
upper limit	-1.4

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	122
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-12.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-26.2
upper limit	1.9

Secondary: Percentage of participants with the B1/NAP1/027 strain of C. difficile at study entry with CDI recurrence

End point title	Percentage of participants with the B1/NAP1/027 strain of C. difficile at study entry with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	24	46	37	36
Units: Percentage of participants				
number (not applicable)	33.3	26.1	10.8	36.1

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC

Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	82
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-10

Confidence interval	
level	95 %
sides	2-sided
lower limit	-30.1
upper limit	10

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	73
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-25.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-43.7
upper limit	-6.1

Secondary: Percentage of participants infected with an epidemic strain of C. difficile (ribotypes 027, 014, 002, 001, 106, and 020) at study entry with CDI recurrence

End point title	Percentage of participants infected with an epidemic strain of C. difficile (ribotypes 027, 014, 002, 001, 106, and 020) at study entry with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	57	108	106	106
Units: Percentage of participants				
number (not applicable)	24.6	23.1	19.8	35.8

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	163
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-11.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-24.9
upper limit	3.9

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	214
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-12.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-24.7
upper limit	-0.5

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description: Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	212
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-16
Confidence interval	
level	95 %
sides	2-sided
lower limit	-27.8
upper limit	-4

Secondary: Percentage of participants with compromised immunity at study entry with CDI recurrence

End point title	Percentage of participants with compromised immunity at study entry with CDI recurrence

End point description:

CDI recurrence is defined as the development of a new episode of diarrhea (3 or more loose stools in 24 or fewer hours) and a positive local or central lab stool test for toxigenic C. difficile following clinical cure of the initial CDI episode. Compromised immunity is defined as follows: an active hematological malignancy (including leukemia, lymphoma, multiple myeloma), an active malignancy requiring recent cytotoxic chemotherapy, receipt of a prior hematopoietic stem cell transplant, receipt of a prior solid organ transplant, asplenia, or neutropenia/pancytopenia due to other conditions. The population analyzed consisted of participants who received infusion of study medication; had a positive local stool test for toxigenic C. difficile; received protocol defined standard of care therapy within 1 day window of the infusion; and complied with Good Clinical Practice.

End point type	Secondary

End point timeframe:

Up to 12 weeks

End point values	MK-3415 + SOC	MK-6072+ SOC	MK-3415A + SOC	Placebo Comparator + SOC
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	55	87	78	92
Units: Percentage of participants				
number (not applicable)	18.2	17.2	11.5	28.3

Statistical analyses

Statistical analysis title	Comparison of Treatment Groups

Statistical analysis description:

Percentage Difference: MK-3415 + SOC - Placebo Comparator + SOC

Comparison groups	MK-3415 + SOC v Placebo Comparator + SOC

Number of subjects included in analysis	147
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-10.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-23.2
upper limit	4.6

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-6072 + SOC - Placebo Comparator + SOC	
Comparison groups	MK-6072+ SOC v Placebo Comparator + SOC
Number of subjects included in analysis	179
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-11
Confidence interval	
level	95 %
sides	2-sided
lower limit	-23.2
upper limit	1.4

Statistical analysis title	Comparison of Treatment Groups
Statistical analysis description:	
Percentage Difference: MK-3415A + SOC - Placebo Comparator + SOC	
Comparison groups	MK-3415A + SOC v Placebo Comparator + SOC
Number of subjects included in analysis	170
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Percentage Difference
Point estimate	-16.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-28.4
upper limit	-4.7

Adverse events

Adverse events information	
Timeframe for reporting adverse events: Up to 90 days	
Assessment type	Systematic
Dictionary used	
Dictionary name	MedDRA
Dictionary version	16.1
Reporting groups	
Reporting group title	MK-3415
Reporting group description: -	
Reporting group title	MK-3415A
Reporting group description: -	
Reporting group title	Placebo
Reporting group description: -	
Reporting group title	MK-6072
Reporting group description: -	

Serious adverse events	MK-3415	MK-3415A	Placebo
Total subjects affected by serious adverse events			
subjects affected / exposed	104 / 235 (44.26%)	94 / 387 (24.29%)	126 / 400 (31.50%)
number of deaths (all causes)	27	20	26
number of deaths resulting from adverse events	0	1	0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acute myeloid leukaemia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Acute myeloid leukaemia recurrent			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Adenocarcinoma of colon			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Bladder transitional cell carcinoma			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Breast cancer			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Breast cancer metastatic			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Burkitt's lymphoma			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Gastric cancer			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Hypergammaglobulinaemia benign monoclonal			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Malignant melanoma			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma in situ			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant neoplasm progression			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Metastases to spine			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Myelodysplastic syndrome			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian cancer metastatic			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancreatic carcinoma			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Squamous cell carcinoma			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Deep vein thrombosis			

subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematoma			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertensive crisis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	2 / 235 (0.85%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypovolaemic shock			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral arterial occlusive disease			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral artery stenosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral artery thrombosis			

subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral ischaemia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Adverse drug reaction			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Asthenia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Death			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 1
General physical health deterioration			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hernia obstructive			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multi-organ failure			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 2
Non-cardiac chest pain			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral swelling			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	2 / 235 (0.85%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sudden cardiac death			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Sudden death			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 2
Systemic inflammatory response syndrome			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 0
Immune system disorders			
Acute graft versus host disease			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute graft versus host disease in intestine			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Graft versus host disease			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Graft versus host disease in gastrointestinal tract			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Heart transplant rejection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspiration			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Asthma			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Atelectasis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chronic obstructive pulmonary disease			
subjects affected / exposed	2 / 235 (0.85%)	2 / 387 (0.52%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Chronic respiratory failure			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Cough			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypoxia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pharyngeal stenosis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleural effusion			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	3 / 400 (0.75%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 2
Pneumonia aspiration			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Pneumothorax			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary cavitation			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Respiratory arrest			
subjects affected / exposed	0 / 235 (0.00%)	2 / 387 (0.52%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	1 / 1	0 / 0
Respiratory disorder			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	2 / 235 (0.85%)	1 / 387 (0.26%)	4 / 400 (1.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 4
deaths causally related to treatment / all	0 / 2	0 / 1	0 / 2
Sputum increased			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psychiatric disorders			
Affective disorder			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aggression			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Alcoholism			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Confusional state			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug dependence			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Mental status changes			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psychogenic seizure			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Alanine aminotransferase increased			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood alkaline phosphatase increased			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemoglobin decreased			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Heart rate increased			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Arteriovenous fistula thrombosis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Cervical vertebral fracture			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Contusion			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fall			
subjects affected / exposed	1 / 235 (0.43%)	3 / 387 (0.78%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gun shot wound			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hip fracture			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal anastomosis complication			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Perinephric collection			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post-traumatic pain			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Procedural vomiting			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Radiation oesophagitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rib fracture			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute myocardial infarction			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Angina pectoris			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina unstable			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	2 / 235 (0.85%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial flutter			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial tachycardia			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bradycardia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bundle branch block left			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 1
Cardiac disorder			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			
subjects affected / exposed	3 / 235 (1.28%)	1 / 387 (0.26%)	4 / 400 (1.00%)
occurrences causally related to treatment / all	0 / 4	0 / 1	0 / 4
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 2
Cardiac failure acute			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure chronic			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			

subjects affected / exposed	2 / 235 (0.85%)	3 / 387 (0.78%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 3	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardio-respiratory arrest			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiopulmonary failure			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 2
Hypertensive heart disease			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intrapericardial thrombosis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Myocardial infarction			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Palpitations			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus arrhythmia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tachycardia			

subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ventricular tachyarrhythmia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ventricular tachycardia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Aphasia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral haemorrhage			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral infarction			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Cerebrovascular accident			
subjects affected / exposed	3 / 235 (1.28%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Convulsion			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depressed level of consciousness			

subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Disturbance in attention			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Encephalopathy			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epilepsy			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Headache			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatic encephalopathy			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolic encephalopathy			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Partial seizures			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Presyncope			

subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transient ischaemic attack			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	2 / 235 (0.85%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Disseminated intravascular coagulation			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	1 / 235 (0.43%)	2 / 387 (0.52%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Iron deficiency anaemia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukocytosis			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neutropenia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancytopenia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Hearing impaired			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Eye disorders			
Diplopia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal distension			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Abdominal hernia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal pain			
subjects affected / exposed	4 / 235 (1.70%)	4 / 387 (1.03%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 4	0 / 4	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal pain lower			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal pain upper			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute abdomen			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ascites			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis ulcerative			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			

subjects affected / exposed	3 / 235 (1.28%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Crohn's disease			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			
subjects affected / exposed	6 / 235 (2.55%)	6 / 387 (1.55%)	6 / 400 (1.50%)
occurrences causally related to treatment / all	0 / 6	0 / 8	0 / 7
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Diarrhoea haemorrhagic			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticular perforation			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulum			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epiploic appendagitis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Faecal incontinence			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastritis alcoholic			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Gastrointestinal inflammation			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematochezia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hernial eventration			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ileus			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Incarcerated umbilical hernia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inflammatory bowel disease			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal dilatation			

subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Intestinal obstruction			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Large intestine perforation			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Melaena			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			
subjects affected / exposed	3 / 235 (1.28%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oesophageal stenosis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancreatitis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Pancreatitis acute			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancreatitis necrotising			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Proctalgia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	0 / 235 (0.00%)	2 / 387 (0.52%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Umbilical hernia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Upper gastrointestinal haemorrhage			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Cholangitis			
subjects affected / exposed	0 / 235 (0.00%)	3 / 387 (0.78%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cholecystitis			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cholecystitis acute			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cholelithiasis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cirrhosis alcoholic			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatic cirrhosis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Hepatorenal syndrome			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Portal vein thrombosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Ecchymosis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Eczema			

subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rash			
subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin ulcer			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			
Bladder dilatation			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematuria			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nephritis autoimmune			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal failure acute			
subjects affected / exposed	3 / 235 (1.28%)	3 / 387 (0.78%)	6 / 400 (1.50%)
occurrences causally related to treatment / all	0 / 3	0 / 3	0 / 6
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 1
Renal failure chronic			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal impairment			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary retention			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Compartment syndrome			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Connective tissue disorder			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Flank pain			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemarthrosis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rhabdomyolysis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal disorder			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			

Abdominal abscess			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abscess limb			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Arthritis bacterial			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bacteraemia			
subjects affected / exposed	1 / 235 (0.43%)	3 / 387 (0.78%)	3 / 400 (0.75%)
occurrences causally related to treatment / all	0 / 1	0 / 3	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Biliary tract infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchitis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Bronchitis viral			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchopneumonia			

subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bursitis infective			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Campylobacter gastroenteritis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Candida infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	3 / 400 (0.75%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	26 / 235 (11.06%)	18 / 387 (4.65%)	26 / 400 (6.50%)
occurrences causally related to treatment / all	0 / 28	0 / 21	0 / 32
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile sepsis			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Cystitis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			

subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Encephalitis viral			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Erysipelas			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gangrene			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gas gangrene			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis			

subjects affected / exposed	0 / 235 (0.00%)	3 / 387 (0.78%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	1 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Graft infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Histoplasmosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypopyon			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infected lymphocele			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Kidney infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lobar pneumonia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			

subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower respiratory tract infection			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Mediastinitis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Osteomyelitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peritonitis			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			
subjects affected / exposed	7 / 235 (2.98%)	5 / 387 (1.29%)	11 / 400 (2.75%)
occurrences causally related to treatment / all	0 / 7	0 / 5	0 / 11
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 4
Pneumonia bacterial			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia haemophilus			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural infection			

subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Postoperative wound infection			
subjects affected / exposed	0 / 235 (0.00%)	2 / 387 (0.52%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pseudomembranous colitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psoas abscess			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Rhinovirus infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Salmonellosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sepsis			

subjects affected / exposed	9 / 235 (3.83%)	3 / 387 (0.78%)	11 / 400 (2.75%)
occurrences causally related to treatment / all	0 / 9	1 / 3	0 / 11
deaths causally related to treatment / all	0 / 7	1 / 1	0 / 3
Septic shock			
subjects affected / exposed	3 / 235 (1.28%)	4 / 387 (1.03%)	4 / 400 (1.00%)
occurrences causally related to treatment / all	0 / 3	0 / 4	0 / 4
deaths causally related to treatment / all	0 / 3	0 / 1	0 / 4
Soft tissue infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Systemic candida			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Systemic mycosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tracheitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tracheobronchitis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Upper respiratory tract infection			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			

subjects affected / exposed	6 / 235 (2.55%)	4 / 387 (1.03%)	5 / 400 (1.25%)
occurrences causally related to treatment / all	0 / 7	0 / 4	0 / 5
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Urosepsis			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	2 / 400 (0.50%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Wound infection			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Zygomycosis			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	1 / 235 (0.43%)	3 / 387 (0.78%)	5 / 400 (1.25%)
occurrences causally related to treatment / all	0 / 1	0 / 3	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Electrolyte imbalance			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	1 / 400 (0.25%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Fluid overload			
subjects affected / exposed	2 / 235 (0.85%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			

subjects affected / exposed	1 / 235 (0.43%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperammonaemia			
subjects affected / exposed	1 / 235 (0.43%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemic hyperosmolar nonketotic syndrome			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			
subjects affected / exposed	0 / 235 (0.00%)	0 / 387 (0.00%)	3 / 400 (0.75%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypoglycaemia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	1 / 1	0 / 0
Hypokalaemia			
subjects affected / exposed	0 / 235 (0.00%)	1 / 387 (0.26%)	0 / 400 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	MK-6072		
Total subjects affected by serious adverse events			
subjects affected / exposed	120 / 390 (30.77%)		
number of deaths (all causes)	31		
number of deaths resulting from adverse events	1		

Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acute myeloid leukaemia			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Acute myeloid leukaemia recurrent			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Adenocarcinoma of colon			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Bladder transitional cell carcinoma			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Breast cancer			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Breast cancer metastatic			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Burkitt's lymphoma			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Colon cancer metastatic			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		

Gastric cancer			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hypergammaglobulinaemia benign monoclonal			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Lung neoplasm malignant			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Malignant melanoma			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Malignant melanoma in situ			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Malignant neoplasm progression			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Metastases to spine			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Myelodysplastic syndrome			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Ovarian cancer metastatic			

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pancreatic carcinoma				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Squamous cell carcinoma				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Vascular disorders				
Deep vein thrombosis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Haematoma				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hypertension				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hypertensive crisis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Hypotension				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hypovolaemic shock				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Peripheral arterial occlusive disease				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Peripheral artery stenosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Peripheral artery thrombosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Peripheral ischaemia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
General disorders and administration site conditions				
Adverse drug reaction				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Asthenia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Chest pain				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Death				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
General physical health deterioration				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hernia obstructive				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Malaise				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Multi-organ failure				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Non-cardiac chest pain				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Peripheral swelling				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pyrexia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Sudden cardiac death				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Sudden death				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Systemic inflammatory response syndrome				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Immune system disorders				
Acute graft versus host disease				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Acute graft versus host disease in intestine				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Graft versus host disease				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Graft versus host disease in gastrointestinal tract				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Heart transplant rejection				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			

Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Aspiration			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Asthma			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Atelectasis			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Chronic obstructive pulmonary disease			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 2		
Chronic respiratory failure			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cough			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hypoxia			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		

Pharyngeal stenosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pleural effusion				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Pneumonia aspiration				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumothorax				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Pulmonary cavitation				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pulmonary embolism				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pulmonary oedema				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Respiratory arrest				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Respiratory disorder				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Respiratory failure				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 2			
Sputum increased				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Psychiatric disorders				
Affective disorder				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Aggression				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Alcoholism				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Confusional state				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Depression				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Drug dependence				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Mental status changes				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Psychogenic seizure				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Suicide attempt				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Investigations				
Alanine aminotransferase increased				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Blood alkaline phosphatase increased				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Blood bilirubin increased				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Haemoglobin decreased				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Heart rate increased				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Platelet count decreased				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Injury, poisoning and procedural complications				
Arteriovenous fistula thrombosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cervical vertebral fracture				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Contusion				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Fall				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gun shot wound				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hip fracture				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Intestinal anastomosis complication				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Perinephric collection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Post-traumatic pain				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Procedural vomiting				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Radiation oesophagitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Rib fracture				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cardiac disorders				
Acute myocardial infarction				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Angina pectoris				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Angina unstable				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Atrial fibrillation				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Atrial flutter				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Atrial tachycardia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bradycardia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bundle branch block left				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cardiac arrest				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Cardiac disorder				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Cardiac failure				

subjects affected / exposed	4 / 390 (1.03%)			
occurrences causally related to treatment / all	0 / 4			
deaths causally related to treatment / all	0 / 1			
Cardiac failure acute				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cardiac failure chronic				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Cardiac failure congestive				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Cardio-respiratory arrest				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Cardiopulmonary failure				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Hypertensive heart disease				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Intrapericardial thrombosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Myocardial infarction				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Palpitations				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Sinus arrhythmia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Tachycardia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Ventricular tachyarrhythmia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Ventricular tachycardia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Nervous system disorders				
Aphasia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cerebral haemorrhage				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	1 / 1			
Cerebral infarction				

subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cerebrovascular accident			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 2		
Convulsion			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Depressed level of consciousness			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Disturbance in attention			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Encephalopathy			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Epilepsy			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Headache			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hepatic encephalopathy			

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Metabolic encephalopathy				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Partial seizures				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Presyncope				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Subarachnoid haemorrhage				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Syncope				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Transient ischaemic attack				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Blood and lymphatic system disorders				
Anaemia				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Disseminated intravascular coagulation				

subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Febrile neutropenia			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Iron deficiency anaemia			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Leukocytosis			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Neutropenia			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Pancytopenia			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Thrombocytopenia			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Ear and labyrinth disorders			
Hearing impaired			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Eye disorders			

Diplopia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gastrointestinal disorders				
Abdominal discomfort				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abdominal distension				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abdominal hernia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abdominal pain				
subjects affected / exposed	3 / 390 (0.77%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 0			
Abdominal pain lower				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abdominal pain upper				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Acute abdomen				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Ascites				

subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Colitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Colitis ulcerative				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Constipation				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Crohn's disease				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Diarrhoea				
subjects affected / exposed	9 / 390 (2.31%)			
occurrences causally related to treatment / all	1 / 10			
deaths causally related to treatment / all	0 / 0			
Diarrhoea haemorrhagic				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Diverticular perforation				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Diverticulum				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Epiploic appendagitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Faecal incontinence				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gastritis alcoholic				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gastrointestinal haemorrhage				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Gastrointestinal inflammation				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Haematochezia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hernial eventration				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Ileus				

subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Incarcerated umbilical hernia			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Inflammatory bowel disease			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Intestinal dilatation			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Intestinal obstruction			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Large intestine perforation			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Melaena			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Nausea			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Oesophageal stenosis			

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pancreatitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pancreatitis acute				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pancreatitis necrotising				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Proctalgia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Rectal haemorrhage				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Small intestinal obstruction				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Umbilical hernia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Upper gastrointestinal haemorrhage				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Vomiting				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hepatobiliary disorders				
Cholangitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Cholecystitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Cholecystitis acute				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cholelithiasis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cirrhosis alcoholic				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hepatic cirrhosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hepatorenal syndrome				

subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Portal vein thrombosis			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Skin and subcutaneous tissue disorders			
Ecchymosis			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Eczema			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Rash			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Skin ulcer			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Renal and urinary disorders			
Bladder dilatation			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Haematuria			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Nephritis autoimmune			

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Renal failure acute				
subjects affected / exposed	5 / 390 (1.28%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 2			
Renal failure chronic				
subjects affected / exposed	3 / 390 (0.77%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 2			
Renal impairment				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Urinary retention				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Musculoskeletal and connective tissue disorders				
Compartment syndrome				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Connective tissue disorder				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Flank pain				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Haemarthrosis				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Rhabdomyolysis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Spinal disorder				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Infections and infestations				
Abdominal abscess				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abdominal infection				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Abscess limb				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Arthritis bacterial				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bacteraemia				
subjects affected / exposed	3 / 390 (0.77%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 0			
Biliary tract infection				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Bronchitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bronchitis viral				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Bronchopneumonia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bursitis infective				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Campylobacter gastroenteritis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Candida infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cellulitis				
subjects affected / exposed	3 / 390 (0.77%)			
occurrences causally related to treatment / all	0 / 4			
deaths causally related to treatment / all	0 / 0			
Clostridium difficile infection				

subjects affected / exposed	10 / 390 (2.56%)		
occurrences causally related to treatment / all	0 / 13		
deaths causally related to treatment / all	0 / 0		
Clostridium difficile sepsis			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cystitis			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Device related infection			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Device related sepsis			
subjects affected / exposed	2 / 390 (0.51%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Diverticulitis			
subjects affected / exposed	1 / 390 (0.26%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Empyema			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Encephalitis viral			
subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Erysipelas			

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gangrene				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Gas gangrene				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gastroenteritis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Gastroenteritis viral				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Graft infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Histoplasmosis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Hypopyon				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Infected lymphocele				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Kidney infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Lobar pneumonia				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 1			
Localised infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Lower respiratory tract infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Mediastinitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Osteomyelitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Peritonitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pneumonia				

subjects affected / exposed	7 / 390 (1.79%)			
occurrences causally related to treatment / all	0 / 7			
deaths causally related to treatment / all	0 / 2			
Pneumonia bacterial				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia haemophilus				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Post procedural infection				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Postoperative wound infection				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pseudomembranous colitis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Psoas abscess				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pyelonephritis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Respiratory tract infection				

subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Rhinovirus infection				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Salmonellosis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Sepsis				
subjects affected / exposed	7 / 390 (1.79%)			
occurrences causally related to treatment / all	1 / 7			
deaths causally related to treatment / all	0 / 3			
Septic shock				
subjects affected / exposed	3 / 390 (0.77%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 3			
Soft tissue infection				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Systemic candida				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Systemic mycosis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Tracheitis				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Tracheobronchitis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Upper respiratory tract infection				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Urinary tract infection				
subjects affected / exposed	6 / 390 (1.54%)			
occurrences causally related to treatment / all	0 / 7			
deaths causally related to treatment / all	0 / 0			
Urosepsis				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Wound infection				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 1			
Zygomycosis				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Metabolism and nutrition disorders				
Dehydration				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Electrolyte imbalance				

subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Failure to thrive				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Fluid overload				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Gout				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hyperammonaemia				
subjects affected / exposed	0 / 390 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hyperglycaemia				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Hyperglycaemic hyperosmolar nonketotic syndrome				
subjects affected / exposed	1 / 390 (0.26%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Hyperkalaemia				
subjects affected / exposed	2 / 390 (0.51%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 1			
Hypoglycaemia				

subjects affected / exposed	0 / 390 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hypokalaemia			
subjects affected / exposed	1 / 390 (0.26%)		
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	MK-3415	MK-3415A	Placebo
Total subjects affected by non-serious adverse events			
subjects affected / exposed	84 / 235 (35.74%)	110 / 387 (28.42%)	103 / 400 (25.75%)
Nervous system disorders			
Headache			
subjects affected / exposed	15 / 235 (6.38%)	22 / 387 (5.68%)	14 / 400 (3.50%)
occurrences (all)	19	23	18
General disorders and administration site conditions			
Fatigue			
subjects affected / exposed	12 / 235 (5.11%)	14 / 387 (3.62%)	5 / 400 (1.25%)
occurrences (all)	13	15	5
Pyrexia			
subjects affected / exposed	14 / 235 (5.96%)	13 / 387 (3.36%)	15 / 400 (3.75%)
occurrences (all)	18	16	16
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	14 / 235 (5.96%)	20 / 387 (5.17%)	24 / 400 (6.00%)
occurrences (all)	15	21	28
Diarrhoea			
subjects affected / exposed	18 / 235 (7.66%)	33 / 387 (8.53%)	27 / 400 (6.75%)
occurrences (all)	23	38	42
Nausea			
subjects affected / exposed	30 / 235 (12.77%)	33 / 387 (8.53%)	30 / 400 (7.50%)
occurrences (all)	33	40	35
Vomiting			

subjects affected / exposed	11 / 235 (4.68%)	15 / 387 (3.88%)	16 / 400 (4.00%)
occurrences (all)	13	18	16
Infections and infestations			
Urinary tract infection			
subjects affected / exposed	15 / 235 (6.38%)	15 / 387 (3.88%)	20 / 400 (5.00%)
occurrences (all)	15	15	21

Non-serious adverse events	MK-6072		
Total subjects affected by non-serious adverse events			
subjects affected / exposed	116 / 390 (29.74%)		
Nervous system disorders			
Headache			
subjects affected / exposed	20 / 390 (5.13%)		
occurrences (all)	23		
General disorders and administration site conditions			
Fatigue			
subjects affected / exposed	7 / 390 (1.79%)		
occurrences (all)	7		
Pyrexia			
subjects affected / exposed	23 / 390 (5.90%)		
occurrences (all)	26		
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	25 / 390 (6.41%)		
occurrences (all)	32		
Diarrhoea			
subjects affected / exposed	25 / 390 (6.41%)		
occurrences (all)	33		
Nausea			
subjects affected / exposed	32 / 390 (8.21%)		
occurrences (all)	38		
Vomiting			
subjects affected / exposed	23 / 390 (5.90%)		
occurrences (all)	29		
Infections and infestations			
Urinary tract infection			

subjects affected / exposed occurrences (all)	18 / 390 (4.62%) 18		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
15 November 2010	Amendment No. 01 was finalized and approved before any participants were enrolled into the study. Major changes included: updates to the statistical analysis plan; modifications to the study procedures; changes to criteria for standard of care antibiotic switching; and changes to the criteria for when to conduct an unscheduled visit.
05 July 2011	Amendment No. 02 was finalized and approved before any participants were enrolled into the study. Major changes included: addition of fidaxomicin as an allowed standard of care antibiotic; revised eligibility criteria to (1) allow initiation of standard of care therapy within a few hours after the study medication infusion, (2) exclude participants with a condition such that they routinely pass loose stool, (3) exclude participants for whom treatment with standard of care therapy was planned for longer than 14 days; removed collection of several biological samples which had been intended for exploratory analyses; reduced study medication infusion duration from 2 hours to 1 hour; updated the statistical analysis plan to address regulatory agency advice.
20 May 2013	Amendment No.03 was implemented after enrollment of subjects had commenced and before database lock and unblinding. Major changes included: increased infusion set filter pore size to 5 micron or smaller from 0.2 microns or smaller; removed the 9-month extended follow-up portion of the study (no participants had been enrolled in the extension prior to this amendment); updated eligibility criteria to exclude participants who had received an experimental C. difficile vaccine or other experimental monoclonal antibody against C. difficile toxin A or B, or participants who planned to receive during the follow-up period fecal transplantation therapy or any other therapies that had been demonstrated to decrease CDI recurrence; modified the definition of the clinical cure endpoint with regard to the duration of standard of care medication: a 14 day regimen was defined as treatment spanning no more than 16 calendar days.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported