



## Clinical trial results:

### **A PHASE 3, RANDOMIZED, DOUBLE-BLIND, PLACEBO-CONTROLLED STUDY TO COMPARE EFFICACY AND SAFETY OF ORAL AZACITIDINE PLUS BEST SUPPORTIVE CARE VERSUS BEST SUPPORTIVE CARE AS MAINTENANCE THERAPY IN SUBJECTS WITH ACUTE MYELOID LEUKEMIA IN COMPLETE REMISSION**

#### **Summary**

EudraCT number	2012-003457-28
Trial protocol	BE CZ PT LT IT ES FI DE AT PL IE GB FR
Global end of trial date	18 June 2024

#### **Results information**

Result version number	v1 (current)
This version publication date	04 July 2025
First version publication date	04 July 2025

#### **Trial information**

##### **Trial identification**

Sponsor protocol code	CC-486-AML-001
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##### **Additional study identifiers**

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

Notes:

##### **Sponsors**

Sponsor organisation name	Bristol-Myers Squibb
Sponsor organisation address	Chaussée de la Hulpe 185, Brussels, Belgium, 1170
Public contact	Global Submission Management, Clinical Trials, Bristol-Myers Squibb International Corporation, mg-gsm-ct@bms.com
Scientific contact	Bristol-Myers Squibb Study Director, Bristol-Myers Squibb, mg-gsm-ct@bms.com

Notes:

##### **Paediatric regulatory details**

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

Notes:

## Results analysis stage

Analysis stage	Final
Date of interim/final analysis	06 August 2024
Is this the analysis of the primary completion data?	No
Global end of trial reached?	Yes
Global end of trial date	18 June 2024
Was the trial ended prematurely?	No

Notes:

## General information about the trial

Main objective of the trial:

The primary objective of the study is to evaluate whether maintenance therapy with oral azacitidine improves OS compared with placebo in subjects with AML, age  $\geq$  55 years, who have achieved first CR or CRi after induction with intensive chemotherapy with or without consolidation chemotherapy.

Protection of trial subjects:

The study was in compliance with the ethical principles derived from the Declaration of Helsinki and in compliance with all International Conference on Harmonization Good Clinical Practice Guidelines. All the local regulatory requirements pertinent to safety of trial subjects were followed.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	24 April 2013
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	Italy: 75
Country: Number of subjects enrolled	Spain: 38
Country: Number of subjects enrolled	United Kingdom: 35
Country: Number of subjects enrolled	Germany: 32
Country: Number of subjects enrolled	Russian Federation: 19
Country: Number of subjects enrolled	Türkiye: 18
Country: Number of subjects enrolled	Portugal: 15
Country: Number of subjects enrolled	Austria: 14
Country: Number of subjects enrolled	Poland: 14
Country: Number of subjects enrolled	France: 13
Country: Number of subjects enrolled	Belgium: 11
Country: Number of subjects enrolled	Czechia: 10
Country: Number of subjects enrolled	Finland: 9
Country: Number of subjects enrolled	Israel: 9
Country: Number of subjects enrolled	Ireland: 1
Country: Number of subjects enrolled	Lithuania: 1
Country: Number of subjects enrolled	United States: 60
Country: Number of subjects enrolled	Canada: 17
Country: Number of subjects enrolled	Mexico: 2

Country: Number of subjects enrolled	Australia: 49
Country: Number of subjects enrolled	Korea, Republic of: 15
Country: Number of subjects enrolled	Taiwan: 8
Country: Number of subjects enrolled	Brazil: 7
Worldwide total number of subjects	472
EEA total number of subjects	233

Notes:

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### **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)	134
From 65 to 84 years	337
85 years and over	1

## Subject disposition

### Recruitment

Recruitment details: -

### Pre-assignment

Screening details:

472 Participants Randomized, 469 participants treated

### Period 1

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

### Arms

Are arms mutually exclusive?	Yes
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<b>Arm title</b>	Oral Azacitidine Plus Best Supportive Care
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Arm description:

Participants received 300 mg azacitidine tablets once a day (QD) for the first 14 days of each 28-day treatment cycle until discontinuation, which includes the following reasons: disease relapse, withdrawal of consent, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, death, lost to follow-up, or protocol violation or until the end of the study.

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

Dosage and administration details:

150 and/or 200mg

<b>Arm title</b>	Placebo Plus Best Supportive Care
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Arm description:

Participants received identically matching placebo tablets QD for the first 14 days of each 28-day treatment cycle until no longer receiving benefit, withdrawal of consent, disease relapse, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, lost to follow-up, or protocol violation or until the end of the study.

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

Dosage and administration details:

placebo

<b>Number of subjects in period 1</b>	<b>Oral Azacitidine Plus Best Supportive Care</b>	<b>Placebo Plus Best Supportive Care</b>
Started	238	234
Received Treatment	236	233
Completed	0	0
Not completed	238	234
Adverse event, serious fatal	2	2
Physician decision	7	-
Disease Relapse	151	181
Withdrew Consent	19	13
Adverse event, non-fatal	31	11
Other Reasons	28	27

## Baseline characteristics

### Reporting groups

Reporting group title	Oral Azacitidine Plus Best Supportive Care
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Reporting group description:

Participants received 300 mg azacitidine tablets once a day (QD) for the first 14 days of each 28-day treatment cycle until discontinuation, which includes the following reasons: disease relapse, withdrawal of consent, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, death, lost to follow-up, or protocol violation or until the end of the study.

Reporting group title	Placebo Plus Best Supportive Care
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Reporting group description:

Participants received identically matching placebo tablets QD for the first 14 days of each 28-day treatment cycle until no longer receiving benefit, withdrawal of consent, disease relapse, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, lost to follow-up, or protocol violation or until the end of the study.

Reporting group values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care	Total
Number of subjects	238	234	472
Age Categorical Units: Participants			
18 to 64 Years	66	68	134
65 to 84 Years	171	166	337
≥ 85 years	1	0	1
Age Continuous Units: Years			
arithmetic mean	67.9	68.0	
standard deviation	± 5.72	± 5.62	-
Sex: Female, Male Units: Participants			
Female	120	107	227
Male	118	127	245
Ethnicity (NIH/OMB) Units: Subjects			
Hispanic or Latino	20	14	34
Not Hispanic or Latino	196	202	398
Unknown or Not Reported	22	18	40
Race/Ethnicity, Customized Units: Subjects			
White	216	197	413
Black or African-American	2	6	8
Asian	6	20	26
Other	12	11	23
Missing	2	0	2
Initial Acute Myeloid Leukemia (AML) Classification			
AML is classified using the WHO classification system based upon a combination of morphology, immunophenotype, genetics, and clinical features. There are several broad groups and include: 1. AML with genetic abnormalities; 2. AML with multilineage dysplasia 3. AML related to previous chemotherapy or radiation 4. Unspecified AML - do not fall into the above groups			
Units: Subjects			

AML with Recurrent Genetic Abnormalities	39	46	85
AML with Myelodysplasia - Related Changes	49	42	91
Therapy-related Myeloid Neoplasms	2	0	2
AML not Otherwise Specified	148	145	293
Missing	0	1	1
<b>Type of Acute Myeloid Leukemia (AML)</b>			
Primary AML is a cancer that originates in the blood and bone marrow. AML affects a group of white blood cells called myeloid cells, which normally develop into the various types of mature blood cells, such as red blood cells, white blood cells and platelets. Secondary acute myeloid leukemia (s-AML) refers to a leukemic process: (1) evolving from prior myelodysplasia (MDS), myeloproliferative disorder (MPN), or aplastic anemia (AA) with or without treatment or (2) as a product of previous exposure to a proven leukemogenic chemotherapeutic agent (therapy-related AML [t-AML]).			
<b>Units: Subjects</b>			
Primary (de novo)	213	216	429
Secondary	25	18	43
<b>Cytogenetic Risk Category at Diagnosis</b>			
Cytogenetic Risk - Intermediate -I is of a normal karyotype; Poor Risk - includes complex karyotypes having 3 or more cytogenetic abnormalities.			
<b>Units: Subjects</b>			
Intermediate	203	203	406
Poor	35	31	66
<b>Eastern Cooperative Oncology Group (ECOG) Performance Status</b>			
ECOG performance status is used to describe a patient's level of functioning in terms of their ability to care for themselves, daily activity, and physical ability (walking, working, etc.). The scale ranges from 0 to 5: 0 = Fully active, no restrictions; 1 = Restricted activity but ambulatory, able to carry out work of a light nature; 2 = Ambulatory and capable of all self-care but unable to carry out work activities; 3 = Capable to only limited self-care, confined to bed or chair more than 50% of waking hours; 4 = Completely disabled, no self-care, confined to bed or chair; 5 = Dead.			
<b>Units: Subjects</b>			
Grade 0	116	111	227
Grade 1	101	106	207
Grade 2	21	15	36
Grade 3	0	2	2

## End points

### End points reporting groups

Reporting group title	Oral Azacitidine Plus Best Supportive Care
Reporting group description: Participants received 300 mg azacitidine tablets once a day (QD) for the first 14 days of each 28-day treatment cycle until discontinuation, which includes the following reasons: disease relapse, withdrawal of consent, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, death, lost to follow-up, or protocol violation or until the end of the study.	
Reporting group title	Placebo Plus Best Supportive Care
Reporting group description: Participants received identically matching placebo tablets QD for the first 14 days of each 28-day treatment cycle until no longer receiving benefit, withdrawal of consent, disease relapse, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, lost to follow-up, or protocol violation or until the end of the study.	

### Primary: Kaplan-Meier (K-M) Estimate for Overall Survival (OS)

End point title	Kaplan-Meier (K-M) Estimate for Overall Survival (OS)
End point description: Overall survival was defined as time from randomization to death from any cause; participants surviving at the end of the follow-up period, or who withdraw consent, or who were lost to follow up were censored at the date last known alive.	
End point type	Primary
End point timeframe: Day 1 (randomization) up to data cut off date of 15 July 2019; median follow-up for OS estimated by the reverse K-M method was 41.2 months for all participants.	

End point values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	238	234		
Units: Months				
median (confidence interval 95%)	24.7 (18.7 to 30.5)	14.8 (11.7 to 17.6)		

### Statistical analyses

Statistical analysis title	Statistical Analysis for OS
Comparison groups	Oral Azacitidine Plus Best Supportive Care v Placebo Plus Best Supportive Care

Number of subjects included in analysis	472
Analysis specification	Pre-specified
Analysis type	
P-value	= 0.0009 <sup>[1]</sup>
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.69
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.55
upper limit	0.86

Notes:

[1] - The p-value is 2-sided from a log-rank test stratified by age, cytogenetic risk category, and received consolidation therapy or not.

### Secondary: Kaplan-Meier Estimate of Relapse Free Survival (RFS)

End point title	Kaplan-Meier Estimate of Relapse Free Survival (RFS)
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End point description:

RFS was defined as the time from the date of randomization to the date of documented relapse or death from any cause, whichever occurred first. Participants who were still alive without documented relapse, or who were lost to follow-up or withdrew consent without documented relapse, were censored at the date of their last bone marrow assessment, prior to receiving any other therapy for AML. Documented relapse was defined as the earliest date of the following: •  $\geq 5\%$  bone marrow blasts (myeloblasts) from Central Pathology report, or • appearance of  $> 0\%$  blasts in the peripheral blood with a later bone marrow confirmation (bone marrow blast [myeloblasts]  $\geq 5\%$ ) within 100 days, or • at least 2 peripheral blasts  $\geq 5\%$  within 30 days.

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

End point values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	238	234		
Units: Months				
median (confidence interval 95%)	10.2 (7.9 to 12.9)	4.8 (4.6 to 6.4)		

### Statistical analyses

Statistical analysis title	Statistical Analysis for RFS
Comparison groups	Oral Azacitidine Plus Best Supportive Care v Placebo Plus Best Supportive Care

Number of subjects included in analysis	472
Analysis specification	Pre-specified
Analysis type	
P-value	< 0.0001 [2]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.65
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.52
upper limit	0.8

Notes:

[2] - The p-value is 2-sided from a log-rank test stratified by age, cytogenetic risk category, and received consolidation therapy or not.

### Secondary: Kaplan-Meier Estimate of Time to Relapse

End point title	Kaplan-Meier Estimate of Time to Relapse
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End point description:

Time to relapse was defined as the interval (in months) from the date of randomization to the date of documented relapse. Estimates of relapse rate were based on the cumulative incidence function from a competing risk analysis with death as a competing risk for relapse from complete remission (CR)/ complete remission with incomplete blood count recovery (CRi). Documented relapse was defined as, the earliest date of the following: •  $\geq 5\%$  bone marrow blasts (myeloblasts) from Central Pathology report, or • appearance of  $> 0\%$  blasts in the peripheral blood with a later bone marrow confirmation (bone marrow blast [myeloblasts]  $\geq 5\%$ ) within 100 days, or • at least 2 peripheral blasts  $\geq 5\%$  within 30 days.

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

End point values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	238	234		
Units: months				
median (confidence interval 95%)	10.2 (8.3 to 13.4)	4.9 (4.6 to 6.4)		

### Statistical analyses

No statistical analyses for this end point

### Secondary: Kaplan-Meier Estimates of Time to Discontinuation from Treatment

End point title	Kaplan-Meier Estimates of Time to Discontinuation from Treatment
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End point description:

Time to discontinuation from treatment was assessed and defined as the interval from the date of

randomization to the date of discontinuation from study drug. Participants who were receiving treatment at the time of study closure were censored at the date of last visit. Estimates of relapse rate were based on the cumulative incidence function from a competing risk analysis with death as a competing risk for relapse from CR/ CRi.

End point type	Secondary
End point timeframe:	
From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months	

<b>End point values</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	238	234		
Units: months				
median (confidence interval 95%)	14.6 (11.3 to 20.1)	6.9 (5.3 to 7.9)		

### Statistical analyses

No statistical analyses for this end point

### Secondary: Number of Participants with Treatment Emergent Adverse Events (TEAEs)

End point title	Number of Participants with Treatment Emergent Adverse Events (TEAEs)
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End point description:

TEAEs include AEs that started between first dose date and 28 days after the last dose of study drug. A serious adverse event (SAE) is: • Death • Life-threatening event • Inpatient hospitalization or prolongation of existing hospitalization • Persistent or significant disability or incapacity • Congenital anomaly or birth defect • Other important medical event The severity of AEs were assessed by the investigator according to the Common Terminology Criteria for Adverse Events (CTCAE) Version 4.0: Grade 1 (Mild): asymptomatic/mild symptoms; clinical or diagnostic observations only; intervention not indicated. Grade 2 (Moderate): minimal, local or noninvasive intervention indicated; limiting age-appropriate activities of daily living. Grade 3: Severe or medically significant but not immediately life-threatening; hospitalization or prolongation of hospitalization indicated; disabling; limiting self care Grade 4: Life-threatening; urgent intervention indicated. Grade 5: Death due to AE.

End point type	Secondary
End point timeframe:	
From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months	

<b>End point values</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	236	233		
Units: Participants				
≥ 1 TEAE	235	233		

≥ 1 TEAE Related to Study Treatment	213	121		
≥ 1 Serious TEAE	110	109		
≥ 1 Treatment Related Serious TEAE	22	5		
≥ 1 Grade 3/4 TEAE	207	201		
≥ 1 Treatment Related Grade 3/4 TEAE	113	55		
≥ 1 TEAE Leading to Death	15	11		
≥ 1 TEAE Leading to Dose Reduction (Red)	37	6		
≥ 1 TEAE Leading to Dose Interruption	107	43		
≥ 1 TEAE Leading to Dose Red and Interruption	25	3		
≥ 1 TEAE Leading to Study Drug Discontinuation	155	170		

## Statistical analyses

No statistical analyses for this end point

### Secondary: Mean Change in the Functional Assessment of Chronic Illness Therapy (FACIT-Fatigue Scale V 4.0) Score from Baseline

End point title	Mean Change in the Functional Assessment of Chronic Illness Therapy (FACIT-Fatigue Scale V 4.0) Score from Baseline
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End point description:

The functional assessment of chronic illness therapy (FACIT-Fatigue Scale V 4.0) is a subscale of the FACIT-F and has been validated in the oncology setting. The FACIT-Fatigue Scale is a short, 13-item, self-administered tool that measures the level of fatigue in an individual during usually daily activities over the past week. The level of fatigue is measured on a 5-point Likert scale (0 = not at all; 4 = very much). The scores range from 0 to 52, with higher scores indicating less fatigue. If there were missing items, but the participant answered at least 50% of the items, then subscores were prorated.

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

End point values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	165	150		
Units: units on a scale				
arithmetic mean (standard deviation)	-3.7 (± 10.92)	-2.5 (± 9.93)		

## Statistical analyses

No statistical analyses for this end point

### Secondary: Mean Change in the European Quality of Life-Five Dimensions-Three Levels (EQ-5D-3L) Score from Baseline

End point title	Mean Change in the European Quality of Life-Five Dimensions-Three Levels (EQ-5D-3L) Score from Baseline
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End point description:

The EQ-5D-3L is a self-administered questionnaire consisting of 5 questions, pertaining to specific health dimensions (ie, mobility, self-care, pain, usual activities, and anxiety/depression) and a health status scale. Each question has 3 levels of severity, corresponding to no problems, moderate problems and severe problems. Canadian population sample weights were used to derive health utility scores. A higher utility score represents a better health state. A clinically meaningful improvement or worsening was defined as at least 0.08 points of improvement or 0.10 points of worsening from baseline, respectively, for the EQ-5D-3L Health Utility Index. The instrument is scored using the United Kingdom (UK) index ranges from -0.594 to 1, where 0 equates to death and 1 equates to full health; -0.594 is considered 'worse than death'.

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

End point values	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	164	150		
Units: Units on a scale				
arithmetic mean (standard deviation)	-0.0416 ( $\pm$ 0.15467)	-0.0152 ( $\pm$ 0.14799)		

## Statistical analyses

No statistical analyses for this end point

### Secondary: Time to Definitive Clinically Meaningful Deterioration for $\geq 2$ Consecutive Visits as Measured Using the EQ-5D HRQoL Scale

End point title	Time to Definitive Clinically Meaningful Deterioration for $\geq 2$ Consecutive Visits as Measured Using the EQ-5D HRQoL Scale
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End point description:

Clinically meaningful deterioration is defined as a decrease of at least 0.10 points from baseline for at least 2 consecutive visits on the EQ-5D Health Utility Index. The EQ-5D-3L is a self-administered questionnaire with 5 questions covering health dimensions like mobility, self-care, pain, usual activities, and anxiety/depression, plus a health status scale. Each question has 3 severity levels: no problems, moderate problems, and severe problems. Canadian population sample weights derive health utility scores, where higher scores indicate better health. Clinically meaningful improvement or worsening is defined as at least 0.08 points of improvement or 0.10 points of worsening from baseline. The EQ-5D-3L uses the UK index, ranging from -0.594 to 1, where 0 equates to death and 1 to full health; -0.594 is considered 'worse than death'.

here "99999" means NA

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 15 July 2019; approximately 74 months

<b>End point values</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	225	217		
Units: Weeks				
median (confidence interval 95%)	99999 (135.1 to 99999)	99999 (122.9 to 99999)		

## Statistical analyses

<b>Statistical analysis title</b>	Statistical Analysis for EQ-5D
Comparison groups	Oral Azacitidine Plus Best Supportive Care v Placebo Plus Best Supportive Care
Number of subjects included in analysis	442
Analysis specification	Pre-specified
Analysis type	
P-value	= 0.7522 <sup>[3]</sup>
Method	Regression, Cox
Parameter estimate	Hazard ratio (HR)
Point estimate	0.9345
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.6136
upper limit	1.4231

Notes:

[3] - Stratification factors: • Age (at induction therapy): 55 to 64 years and ≥ 65 years • Prior history of MDS: yes/no • Cytogenetic risk (at induction therapy): intermediate-risk/poor-risk • Received consolidation therapy following induction: yes/no

## Secondary: Healthcare Resource Utilization (HRU): Rate of Hospital Events Per Person Year

End point title	Healthcare Resource Utilization (HRU): Rate of Hospital Events Per Person Year
End point description:	HRU is defined as any consumption of healthcare resources directly or indirectly related to the treatment of the patient. HRU is a key component to understand treatment costs and budget impact of new treatments from a provider perspective.
End point type	Secondary
End point timeframe:	From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

<b>End point values</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	236	233		
Units: Hospitalizations per person-years				
number (confidence interval 95%)	0.36 (0.31 to 0.42)	0.63 (0.54 to 0.74)		

### Statistical analyses

No statistical analyses for this end point

### Secondary: Healthcare Resource Utilization (HRU): Number of Days Hospitalized Per Person-Year

End point title	Healthcare Resource Utilization (HRU): Number of Days Hospitalized Per Person-Year
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End point description:

HRU is defined as any consumption of healthcare resources directly or indirectly related to the treatment of the patient. HRU is a key component to understand treatment costs and budget impact of new treatments from a provider perspective.

End point type	Secondary
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End point timeframe:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

<b>End point values</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	236	233		
Units: Days per person-years				
number (confidence interval 95%)	6.00 (5.78 to 6.22)	13.13 (12.68 to 13.60)		

### Statistical analyses

No statistical analyses for this end point

## Adverse events

### Adverse events information

Timeframe for reporting adverse events:

From day 1 (randomization) up to data cut off date of 06 August 2024; approximately 135.5 months

Adverse event reporting additional description:

The number at Risk for All-Cause Mortality represents all Randomized Participants. The number at Risk for Serious Adverse Events and Other (Not Including Serious) Adverse Events represents all participants that received at least 1 dose of study medication

Assessment type	Systematic
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### Dictionary used

Dictionary name	MedDRA
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Dictionary version	27
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### Reporting groups

Reporting group title	Oral Azacitidine Plus Best Supportive Care
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Reporting group description:

Participants received 300 mg azacitidine tablets once a day (QD) for the first 14 days of each 28-day treatment cycle until discontinuation, which includes the following reasons: disease relapse, withdrawal of consent, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, death, lost to follow-up, or protocol violation or until the end of the study.

Reporting group title	Placebo Plus Best Supportive Care
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Reporting group description:

Participants received identically matching placebo tablets QD for the first 14 days of each 28-day treatment cycle until no longer receiving benefit, withdrawal of consent, disease relapse, adverse events, participant became eligible for allogeneic bone marrow or stem cell transplantation during the treatment period, lost to follow-up, or protocol violation or until the end of the study.

<b>Serious adverse events</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care	
Total subjects affected by serious adverse events			
subjects affected / exposed	110 / 236 (46.61%)	109 / 233 (46.78%)	
number of deaths (all causes)	171	175	
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Central nervous system leukaemia			
subjects affected / exposed	2 / 236 (0.85%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Basal cell carcinoma			
subjects affected / exposed	4 / 236 (1.69%)	3 / 233 (1.29%)	
occurrences causally related to treatment / all	0 / 5	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	

Acute myeloid leukaemia recurrent subjects affected / exposed	37 / 236 (15.68%)	58 / 233 (24.89%)
occurrences causally related to treatment / all	0 / 37	0 / 58
deaths causally related to treatment / all	0 / 3	0 / 3
Acute myeloid leukaemia subjects affected / exposed	2 / 236 (0.85%)	5 / 233 (2.15%)
occurrences causally related to treatment / all	0 / 2	0 / 5
deaths causally related to treatment / all	0 / 1	0 / 4
Cholangiocarcinoma subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Chloroma subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Endometrial cancer subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gliomatosis cerebri subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Prostate cancer subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Malignant melanoma subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Meningioma		

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Metastases to meninges</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Lung adenocarcinoma</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Squamous cell carcinoma</b>			
subjects affected / exposed	0 / 236 (0.00%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Prostatic adenoma</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Squamous cell carcinoma of lung</b>			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Squamous cell carcinoma of skin</b>			
subjects affected / exposed	2 / 236 (0.85%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Vascular disorders</b>			
<b>Deep vein thrombosis</b>			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Hypertension</b>			

subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypotension			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery stenosis			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombophlebitis			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fatigue			
subjects affected / exposed	2 / 236 (0.85%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	1 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple organ dysfunction syndrome			
subjects affected / exposed	1 / 236 (0.42%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 2	
General physical health deterioration			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Pyrexia			

subjects affected / exposed	6 / 236 (2.54%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 6	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Immune system disorders</b>			
Allergy to vaccine			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Respiratory, thoracic and mediastinal disorders</b>			
Asthma			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Epistaxis			
subjects affected / exposed	0 / 236 (0.00%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonitis			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary embolism			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary haemorrhage			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Respiratory failure subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psychiatric disorders			
Completed suicide subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Confusional state			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
C-reactive protein increased subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alanine aminotransferase increased subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gamma-glutamyltransferase increased			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
General physical condition abnormal subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Troponin increased subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Weight decreased			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Hip fracture			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Foot fracture			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femoral neck fracture			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cataract traumatic			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chemical peritonitis			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fall			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic intracranial haemorrhage			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Subdural haematoma			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal fracture			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post-traumatic pain			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Congenital, familial and genetic disorders			
Hydrocele			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Acute myocardial infarction			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina pectoris			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina unstable			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic valve disease			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Atrial fibrillation			
subjects affected / exposed	3 / 236 (1.27%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure congestive			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiogenic shock			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Coronary artery disease			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stress cardiomyopathy			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Cerebral ischaemia			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral infarction			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral haemorrhage			
subjects affected / exposed	2 / 236 (0.85%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 2	0 / 1	
Central nervous system inflammation			

subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Haemorrhage intracranial</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
<b>Syncope</b>			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Transient ischaemic attack</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Ischaemic cerebral infarction</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Blood and lymphatic system disorders</b>			
<b>Anaemia</b>			
subjects affected / exposed	2 / 236 (0.85%)	3 / 233 (1.29%)	
occurrences causally related to treatment / all	1 / 2	1 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Disseminated intravascular coagulation</b>			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Febrile neutropenia</b>			
subjects affected / exposed	16 / 236 (6.78%)	9 / 233 (3.86%)	
occurrences causally related to treatment / all	5 / 16	2 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Lymphadenitis</b>			

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Neutropenia</b>			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	2 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Thrombocytopenia</b>			
subjects affected / exposed	2 / 236 (0.85%)	3 / 233 (1.29%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Pancytopenia</b>			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Thrombocytosis</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Eye disorders</b>			
<b>Iridocyclitis</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Keratitis</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Ulcerative keratitis</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Gastrointestinal disorders</b>			

Abdominal pain			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colitis			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Crohn's disease			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Constipation			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diarrhoea			
subjects affected / exposed	3 / 236 (1.27%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 4	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dysphagia			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastritis			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	2 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis eosinophilic			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ileus paralytic			

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Gastrointestinal haemorrhage</b>		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Lower gastrointestinal haemorrhage</b>		
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Inguinal hernia</b>		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Melaena</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Nausea</b>		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Neutropenic colitis</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Small intestinal haemorrhage</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Pancreatitis acute</b>		

subjects affected / exposed	1 / 236 (0.42%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	1 / 1	1 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Small intestinal obstruction			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Upper gastrointestinal haemorrhage			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vomiting			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Cholecystitis			
subjects affected / exposed	3 / 236 (1.27%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis acute			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholelithiasis			
subjects affected / exposed	0 / 236 (0.00%)	2 / 233 (0.86%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis chronic			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperbilirubinaemia			

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Skin and subcutaneous tissue disorders</b>			
Acute febrile neutrophilic dermatosis			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Renal and urinary disorders</b>			
Acute kidney injury			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ureterolithiasis			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Musculoskeletal and connective tissue disorders</b>			
Back pain			
subjects affected / exposed	3 / 236 (1.27%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone pain			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral disc protrusion			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Joint effusion			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Osteoporotic fracture			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteoarthritis			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myalgia			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal pain			
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Infections and infestations</b>			
Atypical pneumonia			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacteraemia			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacterial infection			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
COVID-19			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacterial sepsis			

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Cellulitis		
subjects affected / exposed	4 / 236 (1.69%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	1 / 4	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cholecystitis infective		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Clostridium difficile infection		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Cystitis		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Device related infection		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Device related sepsis		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Diverticulitis		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Endophthalmitis		

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis viral		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis salmonella		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis Escherichia coli		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis		
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastrointestinal infection		
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Herpes zoster		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	1 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Influenza		
subjects affected / exposed	3 / 236 (1.27%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Klebsiella sepsis		

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Lung abscess		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumonia aspiration		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0
Pneumonia		
subjects affected / exposed	13 / 236 (5.51%)	7 / 233 (3.00%)
occurrences causally related to treatment / all	8 / 14	0 / 8
deaths causally related to treatment / all	0 / 0	0 / 0
Osteomyelitis		
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Neutropenic sepsis		
subjects affected / exposed	2 / 236 (0.85%)	2 / 233 (0.86%)
occurrences causally related to treatment / all	1 / 2	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 0
Pneumonia fungal		
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pseudomonas infection		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Rectal abscess		

subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Respiratory tract infection</b>		
subjects affected / exposed	0 / 236 (0.00%)	1 / 233 (0.43%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Sepsis</b>		
subjects affected / exposed	5 / 236 (2.12%)	4 / 233 (1.72%)
occurrences causally related to treatment / all	2 / 5	0 / 5
deaths causally related to treatment / all	0 / 2	0 / 0
<b>Septic shock</b>		
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
<b>Staphylococcal infection</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Staphylococcal sepsis</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Superinfection bacterial</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Wound infection</b>		
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
<b>Urinary tract infection bacterial</b>		

subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Urosepsis</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Urinary tract infection</b>			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Metabolism and nutrition disorders</b>			
<b>Decreased appetite</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Dehydration</b>			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Hyperkalaemia</b>			
subjects affected / exposed	2 / 236 (0.85%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Hypocalcaemia</b>			
subjects affected / exposed	1 / 236 (0.42%)	0 / 233 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
<b>Hypokalaemia</b>			
subjects affected / exposed	1 / 236 (0.42%)	1 / 233 (0.43%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

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

<b>Non-serious adverse events</b>	Oral Azacitidine Plus Best Supportive Care	Placebo Plus Best Supportive Care	
Total subjects affected by non-serious adverse events subjects affected / exposed	235 / 236 (99.58%)	224 / 233 (96.14%)	
Neoplasms benign, malignant and unspecified (incl cysts and polyps) Acute myeloid leukaemia recurrent subjects affected / exposed occurrences (all)	115 / 236 (48.73%) 117	126 / 233 (54.08%) 128	
Vascular disorders Hypertension subjects affected / exposed occurrences (all)	19 / 236 (8.05%) 21	16 / 233 (6.87%) 18	
General disorders and administration site conditions Asthenia subjects affected / exposed occurrences (all)  Oedema peripheral subjects affected / exposed occurrences (all)  Fatigue subjects affected / exposed occurrences (all)  Influenza like illness subjects affected / exposed occurrences (all)  Pyrexia subjects affected / exposed occurrences (all)	44 / 236 (18.64%) 60  22 / 236 (9.32%) 26  71 / 236 (30.08%) 104  12 / 236 (5.08%) 19  37 / 236 (15.68%) 48	12 / 233 (5.15%) 13  24 / 233 (10.30%) 33  44 / 233 (18.88%) 56  7 / 233 (3.00%) 10  43 / 233 (18.45%) 62	
Respiratory, thoracic and mediastinal disorders Cough subjects affected / exposed occurrences (all)  Dyspnoea	30 / 236 (12.71%) 47	39 / 233 (16.74%) 47	

subjects affected / exposed occurrences (all)	13 / 236 (5.51%) 16	14 / 233 (6.01%) 20	
Epistaxis subjects affected / exposed occurrences (all)	16 / 236 (6.78%) 20	17 / 233 (7.30%) 21	
Oropharyngeal pain subjects affected / exposed occurrences (all)	15 / 236 (6.36%) 19	19 / 233 (8.15%) 23	
Psychiatric disorders Anxiety subjects affected / exposed occurrences (all)	17 / 236 (7.20%) 17	8 / 233 (3.43%) 10	
Insomnia subjects affected / exposed occurrences (all)	22 / 236 (9.32%) 26	23 / 233 (9.87%) 25	
Investigations Alanine aminotransferase increased subjects affected / exposed occurrences (all)	12 / 236 (5.08%) 15	4 / 233 (1.72%) 7	
Injury, poisoning and procedural complications Fall subjects affected / exposed occurrences (all)	14 / 236 (5.93%) 16	4 / 233 (1.72%) 4	
Nervous system disorders Headache subjects affected / exposed occurrences (all)	25 / 236 (10.59%) 36	26 / 233 (11.16%) 34	
Dizziness subjects affected / exposed occurrences (all)	25 / 236 (10.59%) 32	21 / 233 (9.01%) 22	
Blood and lymphatic system disorders Anaemia subjects affected / exposed occurrences (all)	51 / 236 (21.61%) 77	42 / 233 (18.03%) 59	
Febrile neutropenia			

subjects affected / exposed	14 / 236 (5.93%)	10 / 233 (4.29%)
occurrences (all)	16	12
Thrombocytopenia		
subjects affected / exposed	81 / 236 (34.32%)	62 / 233 (26.61%)
occurrences (all)	185	73
Neutropenia		
subjects affected / exposed	104 / 236 (44.07%)	61 / 233 (26.18%)
occurrences (all)	420	87
Leukopenia		
subjects affected / exposed	27 / 236 (11.44%)	19 / 233 (8.15%)
occurrences (all)	62	28
Gastrointestinal disorders		
Abdominal pain		
subjects affected / exposed	32 / 236 (13.56%)	15 / 233 (6.44%)
occurrences (all)	51	18
Constipation		
subjects affected / exposed	92 / 236 (38.98%)	56 / 233 (24.03%)
occurrences (all)	181	75
Abdominal pain upper		
subjects affected / exposed	23 / 236 (9.75%)	13 / 233 (5.58%)
occurrences (all)	31	15
Flatulence		
subjects affected / exposed	12 / 236 (5.08%)	4 / 233 (1.72%)
occurrences (all)	13	6
Haemorrhoids		
subjects affected / exposed	12 / 236 (5.08%)	4 / 233 (1.72%)
occurrences (all)	17	4
Dyspepsia		
subjects affected / exposed	12 / 236 (5.08%)	5 / 233 (2.15%)
occurrences (all)	12	7
Diarrhoea		
subjects affected / exposed	119 / 236 (50.42%)	50 / 233 (21.46%)
occurrences (all)	465	79
Nausea		
subjects affected / exposed	153 / 236 (64.83%)	54 / 233 (23.18%)
occurrences (all)	368	78

Stomatitis subjects affected / exposed occurrences (all)	8 / 236 (3.39%) 9	12 / 233 (5.15%) 14	
Vomiting subjects affected / exposed occurrences (all)	142 / 236 (60.17%) 308	23 / 233 (9.87%) 25	
Skin and subcutaneous tissue disorders Pruritus subjects affected / exposed occurrences (all)	16 / 236 (6.78%) 21	18 / 233 (7.73%) 18	
Musculoskeletal and connective tissue disorders Back pain subjects affected / exposed occurrences (all)	27 / 236 (11.44%) 39	24 / 233 (10.30%) 35	
Arthralgia subjects affected / exposed occurrences (all)	45 / 236 (19.07%) 55	32 / 233 (13.73%) 44	
Pain in extremity subjects affected / exposed occurrences (all)	28 / 236 (11.86%) 32	12 / 233 (5.15%) 13	
Infections and infestations Influenza subjects affected / exposed occurrences (all)	17 / 236 (7.20%) 19	7 / 233 (3.00%) 10	
Nasopharyngitis subjects affected / exposed occurrences (all)	20 / 236 (8.47%) 36	16 / 233 (6.87%) 20	
Bronchitis subjects affected / exposed occurrences (all)	14 / 236 (5.93%) 18	9 / 233 (3.86%) 10	
Upper respiratory tract infection subjects affected / exposed occurrences (all)	32 / 236 (13.56%) 49	32 / 233 (13.73%) 59	
Rhinitis subjects affected / exposed occurrences (all)	13 / 236 (5.51%) 18	4 / 233 (1.72%) 5	

Oral herpes			
subjects affected / exposed	13 / 236 (5.51%)	6 / 233 (2.58%)	
occurrences (all)	15	10	
Urinary tract infection			
subjects affected / exposed	18 / 236 (7.63%)	13 / 233 (5.58%)	
occurrences (all)	28	16	
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	31 / 236 (13.14%)	15 / 233 (6.44%)	
occurrences (all)	42	16	
Hypokalaemia			
subjects affected / exposed	21 / 236 (8.90%)	20 / 233 (8.58%)	
occurrences (all)	34	22	

## More information

### Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
08 November 2018	The primary purpose of this protocol amendment is to add an extension phase (EP) to allow all subjects who are on treatment with oral azacitidine and demonstrating clinical benefit in this protocol to continue to do so in an extension phase. In addition, all subjects who were discontinued from the treatment phase (irrespective of randomization arm) and continuing in the Follow-up Phase, will be followed for survival for at least another 12 months, until death, withdrawal of consent, study closure or until the subject is lost to follow-up.

Notes:

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### Interruptions (globally)

Were there any global interruptions to the trial? No

### Limitations and caveats

None reported