

**Table S2: Gastrointestinal-Related Adverse Events (AE) in Enrolled Participants per Subject Number, Group and Trial Stage**

Adverse Events (AEs) (n=36)	Subjects (total)	AEs (total)	Single Dose Exposure						7-Day Exposure					
	(n=36) (%)	N / Total AEs (n=# of subjects)	Placebo		UC781: 0.1%		UC781: 0.25%		Placebo		UC781: 0.1%		UC781: 0.25%	
			# of subjects (n=12)(%)	# of AEs (# of subjects)	# of subjects (n=12)(%)	# of AEs (# of subjects)	# of subjects (n=12)(%)	# of AEs (# of subjects)	# of subjects (n=12)(%)	# of AEs (# of subjects)	# of subjects (n=12)(%)	# of AEs (# of subjects)	# of subjects (n=12)(%)	# of AEs (# of subjects)
<b>Subjects with No AEs Reported**</b>	<b>7/36 (19%)</b>	<b>NA</b>												
<b>AE: Grade 1</b>	<b>15/36 (42%)</b>	<b>28 (15 subjects)</b>	<b>2 (17%)</b>	<b>9 (2 subjects)</b>	<b>4(33%)</b>	<b>5 (4 subjects)</b>	<b>4 (33%)</b>	<b>5 (4 subjects)</b>	<b>2 (17%)</b>	<b>4 (2 subjects)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>4 (33%)</b>	<b>4 (4 subjects)</b>
⇒ <i>Related to Product</i>	<b>1/36 (3%)</b>	<b>2 (1 subject)</b>	<b>1 (8%)</b>	<b>2(1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
→GI	<b>1/36 (3%)</b>	<b>2 (1 subject)</b>	<b>1(8%)</b>	<b>2 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
Hematochezia	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
Tenderness	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
→non-GI	0	0	0	0	0	0	0	0	0	0	0	0	0	0
⇒ <i>Possibly Related to Product</i>	<b>3/36 (8%)</b>	<b>6 (3 subjects)</b>	<b>1 (8%)</b>	<b>3 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1(8%)</b>	<b>1 (1 subject)</b>
→GI	<b>2/36 (6%)</b>	<b>2 (2 subjects)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
Pain w/defecation	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
Throbbing - rectum	1/36 (3%)	1 (1 subject)	0	0	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0
→non-GI	<b>1/36 (3%)</b>	<b>4 (2 subjects)</b>	<b>1 (8%)</b>	<b>2 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>
⇒ <i>Probably Related to Product</i>	<b>1/36 (3%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
→GI	<b>1/36 (3%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
Bloating	1/36 (3%)	1 (1 subject)	0	0	1(8%)	1 (1 subject)	0	0	0	0	0	0	0	0
→non-GI	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
⇒ <i>Not related to Product</i>	<b>13/36 (36%)</b>	<b>19 (13 subjects)</b>	<b>2 (17%)</b>	<b>4 (2 subjects)</b>	<b>3 (25%)</b>	<b>3 (3 subjects)</b>	<b>4 (33%)</b>	<b>5 (4 subjects)</b>	<b>2 (17%)</b>	<b>3 (2 subjects)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>3 (25%)</b>	<b>3 (3 subjects)</b>
→GI	<b>3/36 (8%)</b>	<b>5 (3 subjects)</b>	<b>1 (8%)</b>	<b>2 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>2 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>
Acid reflux	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
Cramping	1/36 (3%)	1 (1 subject)	0	0	0	0	1(8%)	1 (1 subject)	0	0	0	0	0	0
Diarrhea	1/36 (3%)	1 (1 subject)	0	0	0	0	1(8%)	1 (1 subject)	0	0	0	0	0	0
Eructation	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
Polyps - Colon	1/36 (3%)	1 (1 subject)	0	0	0	0	0	0	0	0	1(8%)	1 (1 subject)	0	0
→non-GI	<b>12/36 (33%)</b>	<b>14 (12 subjects)</b>	<b>2 (17%)</b>	<b>2 (1 subject)</b>	<b>3 (25%)</b>	<b>3 (3 subjects)</b>	<b>3 (25%)</b>	<b>3 (3 subjects)</b>	<b>2 (17%)</b>	<b>3 (2 subjects)</b>	<b>0</b>	<b>0</b>	<b>3 (25%)</b>	<b>3 (3 subjects)</b>
<b>AE: Grade 2</b>	<b>5/36 (14%)</b>	<b>8 (5 subjects)</b>	<b>1 (8%)</b>	<b>4 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>
⇒ <i>Possibly Related to Product</i>	<b>3/36 (8%)</b>	<b>6 (3 subjects)</b>	<b>1 (8%)</b>	<b>4 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>
→GI	<b>3/36 (8%)</b>	<b>5 (3 subjects)</b>	<b>1 (8%)</b>	<b>3 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>
Cramping	1/36 (3%)	1 (1 subject)	1(8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
Diarrhea	3/36 (8%)	3 (3 subjects)	1 (8%)	1 (1 subject)	0	0	1(8%)	1 (1 subject)	0	0	1 (8%)	1 (1 subject)	0	0
Flatulence	1/36 (3%)	1 (1 subject)	1 (8%)	1 (1 subject)	0	0	0	0	0	0	0	0	0	0
→non-GI	<b>1/36 (3%)</b>	<b>1 (1 subject)</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
⇒ <i>Not related to Product</i>	<b>2/36 (6%)</b>	<b>2 (2 subjects)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>
→GI	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
→non-GI	<b>2/36 (6%)</b>	<b>2 (2 subjects)</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>	<b>0</b>	<b>0</b>	<b>1 (8%)</b>	<b>1 (1 subject)</b>
<b>AE: Grade 3</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
<i>None reported</i>	0	0	0	0	0	0	0	0	0	0	0	0	0	0
<b>AE: Grade 4</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>	<b>0</b>
<i>None reported</i>	0	0	0	0	0	0	0	0	0	0	0	0	0	0

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