

Manuka Honey for Esophagitis

Supplemental Table 1. Number of Patients with an Adverse Event by System Organ Class, Term, and Grade Any Relationship to Protocol Treatment

System Organ Class	Supportive Care (n=48)					Liquid Honey (n=50)					Lozenge Honey (n=50)				
	Grade					Grade					Grade				
Term	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
BLOOD AND LYMPHATIC SYSTEM DISORDERS	5	6	3	1	0	9	5	6	0	0	4	4	6	0	0
Anemia	6	6	2	1	0	10	5	5	0	0	4	6	3	0	0
Blood and lymphatic system disorders - Other	0	1	0	0	0	0	0	0	0	0	0	0	1	0	0
Febrile neutropenia	0	0	1	0	0	0	0	1	0	0	0	0	3	0	0
CARDIAC DISORDERS	1	1	2	0	1	2	2	0	0	0	0	0	1	0	0
Atrial fibrillation	0	1	0	0	0	1	1	0	0	0	0	0	1	0	0
Cardiac arrest	0	0	0	0	1	0	0	0	0	0	0	0	0	0	0
Chest pain - cardiac	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Myocardial infarction	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Palpitations	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Sinus tachycardia	0	0	0	0	0	1	1	0	0	0	0	0	0	0	0
Supraventricular tachycardia	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
EAR AND LABYRINTH DISORDERS	2	0	0	0	0	1	0	0	0	0	1	1	0	0	0
Ear pain	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Tinnitus	2	0	0	0	0	1	0	0	0	0	0	1	0	0	0
Vertigo	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
EYE DISORDERS	0	0	0	0	0	0	1	0	0	0	2	0	0	0	0
Blurred vision	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Dry eye	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Eye disorders - Other	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
GASTROINTESTINAL DISORDERS	7	20	8	0	0	9	28	6	0	0	10	18	5	1	0
Abdominal pain	1	0	0	0	0	0	1	0	0	0	0	1	0	0	0
Colonic perforation	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
Constipation	11	2	0	0	0	13	1	1	0	0	10	0	0	0	0

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System Organ Class Term	Supportive Care (n=48) Grade					Liquid Honey (n=50) Grade					Lozenge Honey (n=50) Grade				
	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
Diarrhea	7	2	0	0	0	9	2	1	0	0	2	1	0	0	0
Dry mouth	3	0	0	0	0	5	0	0	0	0	2	0	0	0	0
Dyspepsia	6	4	0	0	0	4	2	0	0	0	1	1	0	0	0
Dysphagia	6	11	0	0	0	5	13	2	0	0	7	10	0	0	0
Esophageal pain	0	2	0	0	0	3	5	0	0	0	1	1	1	0	0
Esophagitis	4	10	6	0	0	3	15	1	0	0	3	9	3	0	0
Flatulence	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0
Gastritis	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Gastroesophageal reflux disease	0	0	0	0	0	2	3	0	0	0	1	2	0	0	0
Gastrointestinal disorders - Other	0	0	0	0	0	1	2	0	0	0	1	0	0	1	0
Gastrointestinal pain	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Mucositis oral	2	0	0	0	0	1	1	0	0	0	3	1	1	0	0
Nausea	9	7	1	0	0	10	12	1	0	0	7	4	0	0	0
Oral pain	0	0	0	0	0	1	0	0	0	0	1	0	0	0	0
Small intestinal obstruction	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Vomiting	5	2	1	0	0	6	0	0	0	0	4	0	0	0	0
GENERAL DISORDERS AND ADMINISTRATION SITE CONDITIONS	15	12	4	0	0	7	22	0	0	0	11	11	1	0	0
Chills	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Edema limbs	1	1	0	0	0	0	1	0	0	0	0	0	0	0	0
Fatigue	13	12	3	0	0	9	19	0	0	0	11	10	1	0	0
Fever	1	0	0	0	0	2	0	0	0	0	1	0	0	0	0
General disorders and administration site conditions - Other	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Infusion related reaction	0	0	1	0	0	0	1	0	0	0	0	0	0	0	0
Irritability	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Localized edema	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Non-cardiac chest pain	1	0	0	0	0	0	3	0	0	0	0	0	0	0	0
Pain	3	0	0	0	0	5	1	0	0	0	3	1	0	0	0

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	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
INFECTIONS AND INFESTATIONS	0	2	4	0	1	1	6	4	0	0	1	3	2	0	1
Infections and infestations - Other	0	0	0	0	0	1	0	1	0	0	2	0	0	0	0
Lung infection	0	1	2	0	0	0	0	0	0	0	0	0	2	0	0
Mucosal infection	0	0	0	0	0	0	3	0	0	0	0	2	0	0	0
Papulopustular rash	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Sepsis	0	0	0	0	1	0	0	0	0	0	0	0	0	0	1
Sinusitis	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Skin infection	0	0	0	0	0	0	1	1	0	0	0	0	0	0	0
Upper respiratory infection	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Urinary tract infection	0	1	2	0	0	0	1	2	0	0	0	1	0	0	0
Vaginal infection	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0
INJURY, POISONING AND PROCEDURAL COMPLICATIONS	7	4	1	0	0	8	5	0	0	0	4	7	1	0	0
Burn	0	0	0	0	0	2	0	0	0	0	0	0	0	0	0
Dermatitis radiation	7	4	1	0	0	4	3	0	0	0	5	5	1	0	0
Fall	0	0	0	0	0	1	0	0	0	0	0	1	0	0	0
Injury, poisoning and procedural complications - Other	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
Radiation recall reaction (dermatologic)	0	0	0	0	0	1	2	0	0	0	0	0	0	0	0
INVESTIGATIONS	3	4	6	3	0	5	6	9	4	0	2	4	4	7	0
Alanine aminotransferase increased	1	0	0	0	0	1	0	0	0	0	0	3	0	1	0
Alkaline phosphatase increased	2	0	0	0	0	3	0	0	0	0	1	0	0	0	0
Aspartate aminotransferase increased	2	0	0	0	0	0	0	0	0	0	0	2	0	1	0
Creatinine increased	1	1	0	0	0	3	2	0	0	0	1	1	1	0	0
GGT increased	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Lipase increased	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Lymphocyte count decreased	1	1	4	1	0	0	0	4	1	0	0	1	1	0	0
Neutrophil count decreased	0	2	3	2	0	1	3	2	2	0	3	2	2	5	0
Platelet count decreased	1	2	0	2	0	5	2	3	0	0	5	1	3	3	0

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	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
Weight loss	2	2	0	0	0	5	4	0	0	0	2	1	0	0	0
White blood cell decreased	1	2	4	2	0	1	6	6	2	0	1	1	5	3	0
METABOLISM AND NUTRITION DISORDERS	9	7	4	1	0	9	12	4	1	0	3	7	6	0	0
Anorexia	5	5	1	0	0	6	11	0	0	0	0	4	1	0	0
Dehydration	1	5	3	0	0	2	6	2	0	0	1	3	2	0	0
Hyperglycemia	2	0	0	0	0	3	2	2	0	0	0	0	2	0	0
Hyperkalemia	0	0	0	0	0	0	0	1	0	0	1	0	0	0	0
Hypernatremia	0	0	0	0	0	0	1	0	0	0	1	0	0	0	0
Hypoalbuminemia	3	2	0	0	0	5	0	0	0	0	5	0	0	0	0
Hypocalcemia	2	0	0	0	0	2	1	0	0	0	1	0	0	0	0
Hypoglycemia	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Hypokalemia	3	1	0	1	0	5	0	0	1	0	0	2	1	0	0
Hypomagnesemia	2	0	0	0	0	3	0	0	0	0	2	1	0	0	0
Hyponatremia	4	0	0	0	0	5	0	1	0	0	2	0	1	0	0
Hypophosphatemia	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
MUSCULOSKELETAL AND CONNECTIVE TISSUE DISORDERS	4	3	0	0	0	4	5	1	0	0	1	1	1	0	0
Arthralgia	1	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Arthritis	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Back pain	0	2	0	0	0	2	1	1	0	0	0	0	0	0	0
Chest wall pain	1	1	0	0	0	2	2	0	0	0	1	0	0	0	0
Flank pain	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
Muscle weakness lower limb	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
Musculoskeletal and connective tissue disorder - Other	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Myalgia	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Pain in extremity	2	0	0	0	0	0	2	0	0	0	0	1	0	0	0

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	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
NEOPLASMS BENIGN, MALIGNANT AND UNSPECIFIED (INCL CYSTS AND POLYPS)	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0
Tumor pain	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0
NERVOUS SYSTEM DISORDERS	10	5	1	0	0	13	6	0	1	0	8	5	0	0	0
Ataxia	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Concentration impairment	1	0	0	0	0	1	0	0	0	0	1	0	0	0	0
Depressed level of consciousness	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Dizziness	3	0	0	0	0	3	2	0	0	0	2	0	0	0	0
Dysgeusia	4	2	0	0	0	5	3	0	0	0	2	2	0	0	0
Dysphasia	0	0	0	0	0	1	0	0	0	0	1	0	0	0	0
Headache	4	1	0	0	0	3	1	0	0	0	2	1	0	0	0
Memory impairment	1	0	0	0	0	2	0	0	0	0	1	0	0	0	0
Nervous system disorders - Other	0	0	0	0	0	0	0	0	1	0	0	0	0	0	0
Paresthesia	2	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Peripheral sensory neuropathy	4	2	0	0	0	2	1	0	0	0	1	1	0	0	0
Presyncope	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Stroke	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
Tremor	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
PSYCHIATRIC DISORDERS	5	2	0	0	0	4	3	1	0	0	1	3	0	0	0
Agitation	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Anxiety	2	2	0	0	0	2	2	0	0	0	1	1	0	0	0
Confusion	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Depression	1	0	0	0	0	3	1	0	0	0	0	0	0	0	0
Insomnia	5	1	0	0	0	3	1	1	0	0	0	2	0	0	0
RENAL AND URINARY DISORDERS	1	1	1	0	0	0	1	0	0	0	0	0	2	0	0
Acute kidney injury	0	0	1	0	0	0	0	0	0	0	0	0	1	0	0
Chronic kidney disease	0	0	0	0	0	0	0	0	0	0	0	0	1	0	0
Hematuria	0	0	0	0	0	0	0	0	0	0	0	1	0	0	0
Renal and urinary disorders - Other	1	1	0	0	0	0	0	0	0	0	0	1	0	0	0
Urinary retention	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0

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	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
RESPIRATORY, THORACIC AND MEDIASTINAL DISORDERS	11	7	2	0	0	19	8	4	1	0	9	9	2	0	0
Atelectasis	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
Bronchopulmonary hemorrhage	1	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Cough	10	3	0	0	0	15	6	1	0	0	7	4	0	0	0
Dyspnea	4	4	1	0	0	8	7	2	1	0	5	7	2	0	0
Epistaxis	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Hiccups	0	1	0	0	0	2	0	0	0	0	2	1	0	0	0
Hoarseness	3	0	0	0	0	2	0	0	0	0	1	0	0	0	0
Laryngeal inflammation	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Laryngeal mucositis	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Laryngeal stenosis	0	0	0	0	0	0	0	1	0	0	0	0	0	0	0
Nasal congestion	1	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Pneumonitis	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Productive cough	2	0	0	0	0	1	1	0	0	0	0	0	0	0	0
Respiratory, thoracic and mediastinal disorders - Other	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Sinus disorder	0	0	1	0	0	0	0	0	0	0	0	0	0	0	0
Sore throat	2	0	0	0	0	2	0	0	0	0	1	0	0	0	0
Voice alteration	0	0	0	0	0	3	0	0	0	0	2	1	0	0	0
Wheezing	0	0	0	0	0	2	1	0	0	0	1	0	0	0	0
SKIN AND SUBCUTANEOUS TISSUE DISORDERS	8	3	0	0	0	8	4	1	0	0	4	1	1	0	0
Alopecia	1	2	0	0	0	0	1	0	0	0	1	0	0	0	0
Dry skin	5	0	0	0	0	1	0	0	0	0	1	0	0	0	0
Erythema multiforme	1	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Palmar-plantar erythrodysesthesia syndrome	0	0	0	0	0	0	0	0	0	0	1	0	0	0	0
Pruritus	5	0	0	0	0	1	1	0	0	0	1	0	0	0	0
Rash acneiform	0	1	0	0	0	2	1	0	0	0	0	0	0	0	0

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	1	2	3	4	5	1	2	3	4	5	1	2	3	4	5
Rash maculo-papular	2	2	0	0	0	3	0	0	0	0	0	1	0	0	0
Skin and subcutaneous tissue disorders - Other	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Skin hyperpigmentation	1	0	0	0	0	1	0	0	0	0	2	0	0	0	0
Skin ulceration	0	0	0	0	0	0	0	1	0	0	0	0	1	0	0
VASCULAR DISORDERS	0	2	2	1	0	2	5	1	0	0	0	2	0	0	0
Flushing	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Hypotension	0	1	1	0	0	2	1	1	0	0	0	1	0	0	0
Phlebitis	0	0	0	0	0	0	1	0	0	0	0	0	0	0	0
Superficial thrombophlebitis	0	1	0	0	0	0	0	0	0	0	0	0	0	0	0
Thromboembolic event	1	0	1	1	0	0	2	0	0	0	0	1	0	0	0

Includes adverse events where relationship to protocol treatment is missing.

Adverse events were graded with CTCAE version 4.