

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|   | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|---|--------------|-----------------------------|------------------------------|
| Participants in population                        | 86           | 84                          | 84                           |
| <b>Cardiac Disorders</b>                          | <b>13</b>    | <b>13</b>                   | <b>18</b>                    |
| Atrial Fibrillation                               | 1            | 1                           | 3                            |
| Atrial Flutter                                    | 0            | 1                           | 1                            |
| Atrial Hypertrophy                                | 1            | 0                           | 0                            |
| Atrioventricular Block First Degree               | 1            | 1                           | 0                            |
| Atrioventricular Block Second Degree              | 2            | 0                           | 3                            |
| Bradycardia                                       | 1            | 0                           | 0                            |
| Bundle Branch Block Left                          | 1            | 0                           | 0                            |
| Bundle Branch Block Right                         | 1            | 1                           | 0                            |
| Cardiac Disorder                                  | 0            | 0                           | 1                            |
| Cardiac Failure Congestive                        | 1            | 0                           | 0                            |
| Myocardial Infarction                             | 4            | 2                           | 4                            |
| Palpitations                                      | 0            | 2                           | 0                            |
| Sinus Arrhythmia                                  | 1            | 0                           | 0                            |
| Sinus Bradycardia                                 | 2            | 7                           | 8                            |
| Supraventricular Extrasystoles                    | 1            | 1                           | 1                            |
| Supraventricular Tachycardia                      | 0            | 1                           | 0                            |
| Tachycardia                                       | 1            | 0                           | 0                            |
| Ventricular Extrasystoles                         | 0            | 2                           | 1                            |
| Ventricular Hypertrophy                           | 1            | 0                           | 0                            |
| Wolff-Parkinson-White Syndrome                    | 0            | 1                           | 0                            |
| <b>Congenital, Familial and Genetic Disorders</b> | <b>0</b>     | <b>1</b>                    | <b>2</b>                     |
| Ventricular Septal Defect                         | 0            | 1                           | 2                            |
| <b>Ear and Labyrinth Disorders</b>                | <b>1</b>     | <b>2</b>                    | <b>1</b>                     |
| Cerumen Impaction                                 | 0            | 1                           | 0                            |
| Ear Pain  | 1            | 0                           | 0                            |
| Tinnitus  | 0            | 1                           | 0                            |
| Vertigo   | 0            | 1                           | 1                            |
| <b>Eye Disorders</b>                              | <b>4</b>     | <b>2</b>                    | <b>1</b>                     |
| Conjunctival Haemorrhage                          | 0            | 1                           | 0                            |
| Conjunctivitis                                    | 2            | 0                           | 0                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|   | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|---|--------------|-----------------------------|------------------------------|
| <b>Eye Disorders</b>  | <b>4</b>     | <b>2</b>                    | <b>1</b>                     |
| Eye Allergy   | 1            | 0                           | 0                            |
| Eye Pruritus  | 1            | 0                           | 0                            |
| Eye Swelling  | 1            | 0                           | 0                            |
| Glaucoma  | 1            | 0                           | 0                            |
| Vision Blurred  | 0            | 1                           | 1                            |
| <b>Gastrointestinal Disorders</b>                           | <b>17</b>    | <b>15</b>                   | <b>21</b>                    |
| Abdominal Discomfort  | 0            | 0                           | 1                            |
| Abdominal Pain  | 1            | 3                           | 1                            |
| Constipation  | 1            | 0                           | 0                            |
| Diarrhoea   | 9            | 5                           | 4                            |
| Dyspepsia   | 1            | 1                           | 1                            |
| Dysphagia   | 0            | 1                           | 0                            |
| Flatulence  | 1            | 0                           | 0                            |
| Gastrointestinal Haemorrhage                                | 0            | 0                           | 1                            |
| Gastrooesophageal Reflux Disease                            | 1            | 0                           | 0                            |
| Glossitis   | 1            | 0                           | 0                            |
| Hiatus Hernia   | 1            | 0                           | 0                            |
| Nausea  | 3            | 3                           | 6                            |
| Rectal Haemorrhage  | 0            | 1                           | 0                            |
| Salivary Hypersecretion                                     | 0            | 0                           | 4                            |
| Stomach Discomfort  | 0            | 0                           | 1                            |
| Vomiting  | 3            | 3                           | 7                            |
| <b>General Disorders and Administration Site Conditions</b> | <b>21</b>    | <b>47</b>                   | <b>40</b>                    |
| Application Site Bleeding                                   | 0            | 1                           | 0                            |
| Application Site Dermatitis                                 | 5            | 9                           | 7                            |
| Application Site Desquamation                               | 0            | 1                           | 0                            |
| Application Site Discharge                                  | 0            | 0                           | 1                            |
| Application Site Discolouration                             | 0            | 1                           | 0                            |
| Application Site Erythema                                   | 3            | 12                          | 15                           |
| Application Site Induration                                 | 1            | 0                           | 0                            |
| Application Site Irritation                                 | 3            | 9                           | 9                            |
| Application Site Pain                                       | 0            | 0                           | 2                            |
| Application Site Perspiration                               | 0            | 0                           | 2                            |
| Application Site Pruritus                                   | 6            | 22                          | 22                           |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|   | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|---|--------------|-----------------------------|------------------------------|
| <b>General Disorders and Administration Site Conditions</b> | <b>21</b>    | <b>47</b>                   | <b>40</b>                    |
| Application Site Reaction                                   | 1            | 0                           | 1                            |
| Application Site Swelling                                   | 0            | 1                           | 2                            |
| Application Site Urticaria                                  | 0            | 2                           | 1                            |
| Application Site Vesicles                                   | 1            | 4                           | 6                            |
| Application Site Warmth                                     | 0            | 1                           | 0                            |
| Asthenia  | 1            | 0                           | 1                            |
| Chest Discomfort  | 0            | 0                           | 2                            |
| Chest Pain  | 0            | 0                           | 2                            |
| Chills  | 1            | 1                           | 1                            |
| Cyst  | 0            | 1                           | 0                            |
| Fatigue   | 1            | 5                           | 5                            |
| Feeling Abnormal  | 0            | 0                           | 1                            |
| Feeling Cold  | 0            | 0                           | 1                            |
| Inflammation  | 0            | 1                           | 0                            |
| Malaise   | 0            | 1                           | 2                            |
| Oedema  | 0            | 2                           | 0                            |
| Oedema Peripheral   | 2            | 1                           | 2                            |
| Pain  | 0            | 1                           | 1                            |
| Pyrexia   | 2            | 0                           | 1                            |
| Secretion Discharge   | 0            | 1                           | 0                            |
| Sudden Death  | 0            | 1                           | 0                            |
| Swelling  | 0            | 1                           | 0                            |
| Ulcer   | 0            | 1                           | 0                            |
| <b>Hepatobiliary Disorders</b>                              | <b>1</b>     | <b>0</b>                    | <b>0</b>                     |
| Hyperbilirubinaemia   | 1            | 0                           | 0                            |
| <b>Immune System Disorders</b>                              | <b>0</b>     | <b>1</b>                    | <b>1</b>                     |
| Hypersensitivity  | 0            | 1                           | 0                            |
| Seasonal Allergy  | 0            | 0                           | 1                            |
| <b>Infections and Infestations</b>                          | <b>16</b>    | <b>10</b>                   | <b>13</b>                    |
| Bronchitis  | 1            | 0                           | 0                            |
| Cellulitis  | 0            | 1                           | 0                            |
| Cervicitis  | 1            | 0                           | 0                            |
| Cystitis  | 1            | 0                           | 1                            |
| Ear Infection   | 2            | 0                           | 0                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|   | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|---|--------------|-----------------------------|------------------------------|
| <b>Infections and Infestations</b>                    | <b>16</b>    | <b>10</b>                   | <b>13</b>                    |
| Gastroenteritis Viral                                 | 1            | 0                           | 0                            |
| Hordeolum   | 0            | 0                           | 1                            |
| Influenza   | 1            | 1                           | 1                            |
| Localised Infection                                   | 1            | 1                           | 0                            |
| Lower Respiratory Tract Infection                     | 0            | 0                           | 1                            |
| Nasopharyngitis                                       | 2            | 4                           | 6                            |
| Onychomycosis   | 0            | 1                           | 0                            |
| Pneumonia   | 0            | 1                           | 0                            |
| Rhinitis  | 0            | 0                           | 1                            |
| Upper Respiratory Tract Infection                     | 6            | 1                           | 3                            |
| Urinary Tract Infection                               | 2            | 0                           | 1                            |
| Vaginal Mycosis                                       | 1            | 0                           | 0                            |
| Viral Infection                                       | 0            | 1                           | 0                            |
| <b>Injury, Poisoning and Procedural Complications</b> | <b>4</b>     | <b>5</b>                    | <b>5</b>                     |
| Contusion   | 1            | 1                           | 2                            |
| Excoriation   | 2            | 1                           | 1                            |
| Facial Bones Fracture                                 | 0            | 0                           | 1                            |
| Fall  | 1            | 2                           | 1                            |
| Hip Fracture  | 1            | 0                           | 2                            |
| Joint Dislocation                                     | 0            | 1                           | 0                            |
| Skin Laceration                                       | 1            | 2                           | 0                            |
| Wound   | 0            | 1                           | 0                            |
| <b>Investigations</b>                                 | <b>10</b>    | <b>7</b>                    | <b>6</b>                     |
| Biopsy  | 0            | 0                           | 1                            |
| Biopsy Prostate                                       | 0            | 0                           | 1                            |
| Blood Alkaline Phosphatase Increased                  | 1            | 0                           | 0                            |
| Blood Cholesterol Increased                           | 0            | 0                           | 1                            |
| Blood Creatine Phosphokinase Increased                | 1            | 0                           | 0                            |
| Blood Glucose Increased                               | 0            | 1                           | 1                            |
| Blood Urine Present                                   | 1            | 0                           | 0                            |
| Body Temperature Increased                            | 0            | 1                           | 0                            |
| Cystoscopy  | 1            | 0                           | 0                            |
| Electrocardiogram St Segment Depression               | 4            | 1                           | 0                            |
| Electrocardiogram t Wave Amplitude Decreased          | 1            | 1                           | 0                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|  | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|--|--------------|-----------------------------|------------------------------|
| <b>Investigations</b>  | <b>10</b>    | <b>7</b>                    | <b>6</b>                     |
| Electrocardiogram t Wave Inversion   | 2            | 1                           | 1                            |
| Heart Rate Increased   | 1            | 0                           | 0                            |
| Heart Rate Irregular   | 1            | 0                           | 0                            |
| Nasal Mucosa Biopsy  | 0            | 1                           | 0                            |
| Neutrophil Count Increased   | 0            | 1                           | 0                            |
| Urine Analysis Abnormal  | 0            | 1                           | 0                            |
| Weight Decreased   | 0            | 0                           | 1                            |
| White Blood Cell Count Increased   | 0            | 1                           | 0                            |
| <b>Metabolism and Nutrition Disorders</b>                                  | <b>6</b>     | <b>1</b>                    | <b>3</b>                     |
| Decreased Appetite   | 1            | 0                           | 1                            |
| Dehydration  | 1            | 0                           | 0                            |
| Diabetes Mellitus  | 1            | 0                           | 0                            |
| Food Craving   | 1            | 1                           | 0                            |
| Hypercholesterolaemia  | 0            | 0                           | 1                            |
| Hyponatraemia  | 1            | 0                           | 0                            |
| Increased Appetite   | 1            | 0                           | 1                            |
| <b>Musculoskeletal and Connective Tissue Disorders</b>                     | <b>5</b>     | <b>7</b>                    | <b>8</b>                     |
| Arthralgia   | 1            | 2                           | 1                            |
| Arthritis  | 1            | 0                           | 1                            |
| Back Pain  | 1            | 1                           | 3                            |
| Flank Pain   | 0            | 0                           | 2                            |
| Muscle Spasms  | 0            | 1                           | 1                            |
| Muscular Weakness  | 0            | 1                           | 0                            |
| Myalgia  | 0            | 0                           | 1                            |
| Pain in Extremity  | 1            | 0                           | 0                            |
| Shoulder Pain  | 1            | 2                           | 0                            |
| <b>Neoplasms Benign, Malignant and Unspecified (Incl Cysts and Polyps)</b> | <b>0</b>     | <b>2</b>                    | <b>1</b>                     |
| Colon Cancer   | 0            | 1                           | 0                            |
| Malignant Fibrous Histiocytoma   | 0            | 1                           | 0                            |
| Prostate Cancer  | 0            | 0                           | 1                            |
| <b>Nervous System Disorders</b>  | <b>12</b>    | <b>20</b>                   | <b>27</b>                    |
| Amnesia  | 0            | 0                           | 1                            |
| Balance Disorder   | 0            | 1                           | 0                            |
| Burning Sensation  | 0            | 0                           | 2                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|  | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|--|--------------|-----------------------------|------------------------------|
| <b>Nervous System Disorders</b>                | <b>12</b>    | <b>20</b>                   | <b>27</b>                    |
| Cognitive Disorder                             | 0            | 0                           | 1                            |
| Complex Partial Seizures                       | 0            | 1                           | 0                            |
| Coordination Abnormal                          | 0            | 1                           | 0                            |
| Dizziness                                      | 2            | 8                           | 12                           |
| Headache                                       | 7            | 3                           | 6                            |
| Hemianopia Homonymous                          | 0            | 1                           | 0                            |
| Hypersomnia                                    | 0            | 0                           | 1                            |
| Lethargy                                       | 0            | 1                           | 1                            |
| Paraesthesia                                   | 0            | 0                           | 1                            |
| Paraesthesia Oral                              | 0            | 1                           | 0                            |
| Parkinson's Disease                            | 1            | 0                           | 0                            |
| Parosmia                                       | 0            | 0                           | 1                            |
| Partial Seizures with Secondary Generalisation | 0            | 0                           | 1                            |
| Psychomotor Hyperactivity                      | 1            | 0                           | 0                            |
| Somnolence                                     | 2            | 3                           | 1                            |
| Stupor   | 0            | 1                           | 0                            |
| Syncope  | 0            | 4                           | 3                            |
| Syncope Vasovagal                              | 0            | 0                           | 1                            |
| Transient Ischaemic Attack                     | 0            | 2                           | 1                            |
| <b>Psychiatric Disorders</b>                   | <b>10</b>    | <b>10</b>                   | <b>9</b>                     |
| Agitation                                      | 2            | 2                           | 1                            |
| Anxiety  | 1            | 3                           | 0                            |
| Completed Suicide                              | 1            | 0                           | 0                            |
| Confusional State                              | 2            | 3                           | 1                            |
| Delirium                                       | 0            | 0                           | 1                            |
| Delusion                                       | 1            | 0                           | 1                            |
| Depressed Mood                                 | 0            | 1                           | 1                            |
| Disorientation                                 | 1            | 0                           | 0                            |
| Hallucination                                  | 0            | 0                           | 1                            |
| Hallucination, Visual                          | 0            | 0                           | 1                            |
| Insomnia                                       | 2            | 0                           | 2                            |
| Irritability                                   | 1            | 1                           | 0                            |
| Libido Decreased                               | 0            | 0                           | 1                            |
| Listless                                       | 0            | 0                           | 1                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|  | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|--|--------------|-----------------------------|------------------------------|
| <b>Psychiatric Disorders</b>                           | <b>10</b>    | <b>10</b>                   | <b>9</b>                     |
| Nightmare  | 0            | 0                           | 1                            |
| Restlessness   | 0            | 1                           | 0                            |
| <b>Renal and Urinary Disorders</b>                     | <b>4</b>     | <b>4</b>                    | <b>3</b>                     |
| Calculus Urethral                                      | 0            | 0                           | 1                            |
| Dysuria  | 1            | 1                           | 0                            |
| Enuresis   | 0            | 1                           | 0                            |
| Incontinence   | 0            | 1                           | 0                            |
| Micturition Urgency                                    | 1            | 1                           | 1                            |
| Nephrolithiasis  | 1            | 0                           | 1                            |
| Pollakiuria  | 1            | 0                           | 0                            |
| <b>Reproductive System and Breast Disorders</b>        | <b>2</b>     | <b>0</b>                    | <b>1</b>                     |
| Benign Prostatic Hyperplasia                           | 1            | 0                           | 1                            |
| Pelvic Pain  | 1            | 0                           | 0                            |
| <b>Respiratory, Thoracic and Mediastinal Disorders</b> | <b>10</b>    | <b>10</b>                   | <b>10</b>                    |
| Allergic Granulomatous Angiitis                        | 0            | 0                           | 1                            |
| Cough  | 3            | 6                           | 5                            |
| Dysphonia  | 0            | 1                           | 0                            |
| Dyspnoea   | 1            | 1                           | 1                            |
| Emphysema  | 1            | 0                           | 0                            |
| Epistaxis  | 0            | 1                           | 2                            |
| Haemoptysis  | 1            | 0                           | 0                            |
| Nasal Congestion                                       | 3            | 1                           | 3                            |
| Pharyngeal Erythema                                    | 0            | 0                           | 1                            |
| Pharyngolaryngeal Pain                                 | 0            | 1                           | 1                            |
| Postnasal Drip   | 1            | 0                           | 0                            |
| Productive Cough                                       | 0            | 0                           | 1                            |
| Rales  | 1            | 0                           | 0                            |
| Respiratory Tract Congestion                           | 0            | 0                           | 1                            |
| Rhinorrhoea  | 0            | 1                           | 1                            |
| <b>Skin and Subcutaneous Tissue Disorders</b>          | <b>21</b>    | <b>42</b>                   | <b>42</b>                    |
| Actinic Keratosis                                      | 0            | 0                           | 1                            |
| Alopecia   | 1            | 0                           | 0                            |
| Blister  | 0            | 5                           | 1                            |
| Cold Sweat   | 1            | 0                           | 0                            |

Analysis of Participants With Specific Adverse Events  
(Safety Analysis Population)

|  | Placebo<br>n | Xanomeline Low<br>Dose<br>n | Xanomeline High<br>Dose<br>n |
|--|--------------|-----------------------------|------------------------------|
| <b>Skin and Subcutaneous Tissue Disorders</b>                              | <b>21</b>    | <b>42</b>                   | <b>42</b>                    |
| Dermatitis Atopic  | 1            | 0                           | 0                            |
| Dermatitis Contact   | 0            | 1                           | 0                            |
| Drug Eruption  | 1            | 0                           | 0                            |
| Erythema   | 9            | 15                          | 14                           |
| Hyperhidrosis  | 2            | 4                           | 8                            |
| Pruritus   | 8            | 23                          | 26                           |
| Pruritus Generalised   | 0            | 1                           | 1                            |
| Rash   | 5            | 13                          | 11                           |
| Rash Erythematous  | 0            | 2                           | 0                            |
| Rash Maculo-Papular  | 0            | 0                           | 1                            |
| Rash Papular   | 0            | 0                           | 1                            |
| Rash Pruritic  | 0            | 1                           | 2                            |
| Skin Exfoliation   | 0            | 1                           | 0                            |
| Skin Irritation  | 3            | 6                           | 5                            |
| Skin Odour Abnormal  | 0            | 0                           | 1                            |
| Skin Ulcer   | 1            | 0                           | 0                            |
| Urticaria  | 0            | 1                           | 1                            |
| <b>Social Circumstances</b>  | <b>0</b>     | <b>0</b>                    | <b>1</b>                     |
| Alcohol Use  | 0            | 0                           | 1                            |
| <b>Surgical and Medical Procedures</b>                                     | <b>2</b>     | <b>1</b>                    | <b>2</b>                     |
| Acrochordon Excision   | 0            | 0                           | 1                            |
| Cataract Operation   | 1            | 1                           | 0                            |
| Eye Laser Surgery  | 1            | 0                           | 0                            |
| Skin Lesion Excision   | 0            | 0                           | 1                            |
| <b>Vascular Disorders</b>  | <b>3</b>     | <b>3</b>                    | <b>2</b>                     |
| Hot Flush  | 0            | 1                           | 0                            |
| Hypertension   | 1            | 1                           | 1                            |
| Hypotension  | 2            | 1                           | 0                            |
| Orthostatic Hypotension  | 1            | 0                           | 0                            |
| Wound Haemorrhage  | 0            | 0                           | 1                            |
| Every subject is counted a single time for each applicable row and column. |              |                             |                              |