| Study title | | | |
| --- | --- | --- | --- |
| 1.2.3 Summary of Adverse Events | | | |
| SAF | | | |
| System Organ Class    Preferred Term | Xan High (n = 436) | Xan Low (n = 454) | Placebo (n = 301) |
| Any Adverse Event | 436 (100%) | 454 (100%) | 301 (100%) |
| CARDIAC DISORDERS | 30 (100%) | 34 (100%) | 27 (100%) |
| ATRIAL FIBRILLATION | 3 (10%) | 3 (8.8%) | 1 (3.7%) |
| ATRIAL FLUTTER | 2 (6.7%) | 1 (2.9%) |  |
| ATRIAL HYPERTROPHY |  |  | 2 (7.4%) |
| ATRIOVENTRICULAR BLOCK FIRST DEGREE |  | 1 (2.9%) | 1 (3.7%) |
| ATRIOVENTRICULAR BLOCK SECOND DEGREE | 2 (6.7%) | 2 (5.9%) | 2 (7.4%) |
| BRADYCARDIA |  |  | 4 (14.8%) |
| BUNDLE BRANCH BLOCK LEFT |  |  | 1 (3.7%) |
| BUNDLE BRANCH BLOCK RIGHT |  | 1 (2.9%) | 2 (7.4%) |
| CARDIAC DISORDER | 1 (3.3%) |  |  |
| CARDIAC FAILURE CONGESTIVE |  |  | 1 (3.7%) |
| MYOCARDIAL INFARCTION | 8 (26.7%) | 4 (11.8%) | 4 (14.8%) |
| PALPITATIONS |  | 2 (5.9%) |  |
| SINUS ARRHYTHMIA |  |  | 2 (7.4%) |
| SINUS BRADYCARDIA | 12 (40%) | 10 (29.4%) | 2 (7.4%) |
| SUPRAVENTRICULAR EXTRASYSTOLES | 1 (3.3%) | 2 (5.9%) | 2 (7.4%) |
| SUPRAVENTRICULAR TACHYCARDIA |  | 2 (5.9%) |  |
| TACHYCARDIA |  |  | 2 (7.4%) |
| VENTRICULAR EXTRASYSTOLES | 1 (3.3%) | 4 (11.8%) |  |
| VENTRICULAR HYPERTROPHY |  |  | 1 (3.7%) |
| WOLFF-PARKINSON-WHITE SYNDROME |  | 2 (5.9%) |  |
| CONGENITAL, FAMILIAL AND GENETIC DISORDERS | 2 (100%) | 1 (100%) |  |
| VENTRICULAR SEPTAL DEFECT | 2 (100%) | 1 (100%) |  |
| EAR AND LABYRINTH DISORDERS | 1 (100%) | 3 (100%) | 2 (100%) |
| CERUMEN IMPACTION |  | 1 (33.3%) |  |
| EAR PAIN |  |  | 2 (100%) |
| TINNITUS |  | 1 (33.3%) |  |
| VERTIGO | 1 (100%) | 1 (33.3%) |  |
| EYE DISORDERS | 2 (100%) | 2 (100%) | 8 (100%) |
| CONJUNCTIVAL HAEMORRHAGE |  | 1 (50%) |  |
| CONJUNCTIVITIS |  |  | 4 (50%) |
| EYE ALLERGY |  |  | 1 (12.5%) |
| EYE PRURITUS |  |  | 1 (12.5%) |
| EYE SWELLING |  |  | 1 (12.5%) |
| GLAUCOMA |  |  | 1 (12.5%) |
| VISION BLURRED | 2 (100%) | 1 (50%) |  |
| GASTROINTESTINAL DISORDERS | 35 (100%) | 26 (100%) | 26 (100%) |
| ABDOMINAL DISCOMFORT | 1 (2.9%) |  |  |
| ABDOMINAL PAIN | 2 (5.7%) | 3 (11.5%) | 1 (3.8%) |
| CONSTIPATION |  |  | 1 (3.8%) |
| DIARRHOEA | 3 (8.6%) | 8 (30.8%) | 10 (38.5%) |
| DYSPEPSIA | 1 (2.9%) | 2 (7.7%) | 2 (7.7%) |
| DYSPHAGIA |  | 1 (3.8%) |  |
| FLATULENCE |  |  | 2 (7.7%) |
| GASTROINTESTINAL HAEMORRHAGE | 1 (2.9%) |  |  |
| GASTROOESOPHAGEAL REFLUX DISEASE |  |  | 1 (3.8%) |
| GLOSSITIS |  |  | 1 (3.8%) |
| HIATUS HERNIA |  |  | 2 (7.7%) |
| NAUSEA | 13 (37.1%) | 5 (19.2%) | 3 (11.5%) |
| RECTAL HAEMORRHAGE |  | 2 (7.7%) |  |
| SALIVARY HYPERSECRETION | 5 (14.3%) |  |  |
| STOMACH DISCOMFORT | 1 (2.9%) |  |  |
| VOMITING | 8 (22.9%) | 5 (19.2%) | 3 (11.5%) |
| GENERAL DISORDERS AND ADMINISTRATION SITE CONDITIONS | 118 (100%) | 126 (100%) | 48 (100%) |
| APPLICATION SITE BLEEDING |  | 1 (0.8%) |  |
| APPLICATION SITE DERMATITIS | 12 (10.2%) | 15 (11.9%) | 9 (18.8%) |
| APPLICATION SITE DESQUAMATION |  | 1 (0.8%) |  |
| APPLICATION SITE DISCHARGE | 1 (0.8%) |  |  |
| APPLICATION SITE DISCOLOURATION |  | 1 (0.8%) |  |
| APPLICATION SITE ERYTHEMA | 22 (18.6%) | 21 (16.7%) | 3 (6.2%) |
| APPLICATION SITE INDURATION |  |  | 1 (2.1%) |
| APPLICATION SITE IRRITATION | 16 (13.6%) | 18 (14.3%) | 7 (14.6%) |
| APPLICATION SITE PAIN | 2 (1.7%) |  |  |
| APPLICATION SITE PERSPIRATION | 3 (2.5%) |  |  |
| APPLICATION SITE PRURITUS | 34 (28.8%) | 34 (27%) | 10 (20.8%) |
| APPLICATION SITE REACTION | 1 (0.8%) |  | 2 (4.2%) |
| APPLICATION SITE SWELLING | 3 (2.5%) | 1 (0.8%) |  |
| APPLICATION SITE URTICARIA | 1 (0.8%) | 2 (1.6%) |  |
| APPLICATION SITE VESICLES | 5 (4.2%) | 6 (4.8%) | 2 (4.2%) |
| APPLICATION SITE WARMTH |  | 2 (1.6%) |  |
| ASTHENIA |  | 1 (0.8%) | 2 (4.2%) |
| CHEST DISCOMFORT | 1 (0.8%) | 1 (0.8%) |  |
| CHEST PAIN | 2 (1.7%) |  |  |
| CHILLS | 1 (0.8%) | 2 (1.6%) | 3 (6.2%) |
| CYST |  | 1 (0.8%) |  |
| FATIGUE | 5 (4.2%) | 5 (4%) | 2 (4.2%) |
| FEELING ABNORMAL | 1 (0.8%) |  |  |
| FEELING COLD | 1 (0.8%) |  |  |
| INFLAMMATION |  | 1 (0.8%) |  |
| MALAISE | 3 (2.5%) | 2 (1.6%) |  |
| OEDEMA |  | 2 (1.6%) |  |
| OEDEMA PERIPHERAL | 3 (2.5%) | 1 (0.8%) | 5 (10.4%) |
| PAIN | 1 (0.8%) | 2 (1.6%) |  |
| PYREXIA |  | 1 (0.8%) | 2 (4.2%) |
| SECRETION DISCHARGE |  | 2 (1.6%) |  |
| SUDDEN DEATH |  | 1 (0.8%) |  |
| SWELLING |  | 1 (0.8%) |  |
| ULCER |  | 1 (0.8%) |  |
| HEPATOBILIARY DISORDERS |  |  | 1 (100%) |
| HYPERBILIRUBINAEMIA |  |  | 1 (100%) |
| IMMUNE SYSTEM DISORDERS | 1 (100%) | 2 (100%) |  |
| HYPERSENSITIVITY |  | 2 (100%) |  |
| SEASONAL ALLERGY | 1 (100%) |  |  |
| INFECTIONS AND INFESTATIONS | 20 (100%) | 18 (100%) | 35 (100%) |
| BRONCHITIS |  |  | 1 (2.9%) |
| CELLULITIS |  | 1 (5.6%) |  |
| CERVICITIS |  |  | 2 (5.7%) |
| CYSTITIS | 1 (5%) |  | 1 (2.9%) |
| EAR INFECTION |  |  | 4 (11.4%) |
| GASTROENTERITIS VIRAL |  |  | 1 (2.9%) |
| HORDEOLUM | 1 (5%) |  |  |
| INFLUENZA | 1 (5%) | 1 (5.6%) | 2 (5.7%) |
| LOCALISED INFECTION |  | 1 (5.6%) | 2 (5.7%) |
| LOWER RESPIRATORY TRACT INFECTION | 2 (10%) |  |  |
| NASOPHARYNGITIS | 8 (40%) | 9 (50%) | 4 (11.4%) |
| ONYCHOMYCOSIS |  | 1 (5.6%) |  |
| PNEUMONIA |  | 2 (11.1%) |  |
| RHINITIS | 1 (5%) |  |  |
| UPPER RESPIRATORY TRACT INFECTION | 5 (25%) | 2 (11.1%) | 12 (34.3%) |
| URINARY TRACT INFECTION | 1 (5%) |  | 4 (11.4%) |
| VAGINAL MYCOSIS |  |  | 2 (5.7%) |
| VIRAL INFECTION |  | 1 (5.6%) |  |
| INJURY, POISONING AND PROCEDURAL COMPLICATIONS | 8 (100%) | 12 (100%) | 9 (100%) |
| CONTUSION | 3 (37.5%) | 3 (25%) | 1 (11.1%) |
| EXCORIATION | 1 (12.5%) | 2 (16.7%) | 3 (33.3%) |
| FACIAL BONES FRACTURE | 1 (12.5%) |  |  |
| FALL | 1 (12.5%) | 2 (16.7%) | 2 (22.2%) |
| HIP FRACTURE | 2 (25%) |  | 2 (22.2%) |
| JOINT DISLOCATION |  | 1 (8.3%) |  |
| SKIN LACERATION |  | 2 (16.7%) | 1 (11.1%) |
| WOUND |  | 2 (16.7%) |  |
| INVESTIGATIONS | 6 (100%) | 15 (100%) | 19 (100%) |
| BIOPSY | 1 (16.7%) |  |  |
| BIOPSY PROSTATE | 1 (16.7%) |  |  |
| BLOOD ALKALINE PHOSPHATASE INCREASED |  |  | 1 (5.3%) |
| BLOOD CHOLESTEROL INCREASED | 1 (16.7%) |  |  |
| BLOOD CREATINE PHOSPHOKINASE INCREASED |  |  | 2 (10.5%) |
| BLOOD GLUCOSE INCREASED | 2 (33.3%) | 1 (6.7%) |  |
| BLOOD URINE PRESENT |  |  | 1 (5.3%) |
| BODY TEMPERATURE INCREASED |  | 1 (6.7%) |  |
| CYSTOSCOPY |  |  | 1 (5.3%) |
| ELECTROCARDIOGRAM ST SEGMENT DEPRESSION |  | 2 (13.3%) | 4 (21.1%) |
| ELECTROCARDIOGRAM T WAVE AMPLITUDE DECREASED |  | 1 (6.7%) | 1 (5.3%) |
| ELECTROCARDIOGRAM T WAVE INVERSION | 1 (16.7%) | 1 (6.7%) | 3 (15.8%) |
| HEART RATE INCREASED |  |  | 2 (10.5%) |
| HEART RATE IRREGULAR |  |  | 4 (21.1%) |
| NASAL MUCOSA BIOPSY |  | 1 (6.7%) |  |
| NEUTROPHIL COUNT INCREASED |  | 2 (13.3%) |  |
| URINE ANALYSIS ABNORMAL |  | 2 (13.3%) |  |
| WEIGHT DECREASED |  | 2 (13.3%) |  |
| WHITE BLOOD CELL COUNT INCREASED |  | 2 (13.3%) |  |
| METABOLISM AND NUTRITION DISORDERS | 5 (100%) | 1 (100%) | 8 (100%) |
| DECREASED APPETITE | 2 (40%) |  | 2 (25%) |
| DEHYDRATION |  |  | 1 (12.5%) |
| DIABETES MELLITUS |  |  | 1 (12.5%) |
| FOOD CRAVING |  | 1 (100%) | 1 (12.5%) |
| HYPERCHOLESTEROLAEMIA | 1 (20%) |  |  |
| HYPONATRAEMIA |  |  | 1 (12.5%) |
| INCREASED APPETITE | 2 (40%) |  | 2 (25%) |
| MUSCULOSKELETAL AND CONNECTIVE TISSUE DISORDERS | 11 (100%) | 10 (100%) | 8 (100%) |
| ARTHRALGIA | 1 (9.1%) | 4 (40%) | 1 (12.5%) |
| ARTHRITIS | 1 (9.1%) |  | 2 (25%) |
| BACK PAIN | 4 (36.4%) | 1 (10%) | 2 (25%) |
| FLANK PAIN | 2 (18.2%) |  |  |
| MUSCLE SPASMS | 2 (18.2%) | 1 (10%) |  |
| MUSCULAR WEAKNESS |  | 2 (20%) |  |
| MYALGIA | 1 (9.1%) |  |  |
| PAIN IN EXTREMITY |  |  | 1 (12.5%) |
| SHOULDER PAIN |  | 2 (20%) | 2 (25%) |
| NEOPLASMS BENIGN, MALIGNANT AND UNSPECIFIED (INCL CYSTS AND POLYPS) | 1 (100%) | 3 (100%) |  |
| COLON CANCER |  | 1 (33.3%) |  |
| MALIGNANT FIBROUS HISTIOCYTOMA |  | 2 (66.7%) |  |
| PROSTATE CANCER | 1 (100%) |  |  |
| NERVOUS SYSTEM DISORDERS | 43 (100%) | 42 (100%) | 16 (100%) |
| AMNESIA | 2 (4.7%) |  |  |
| BALANCE DISORDER |  | 3 (7.1%) |  |
| BURNING SENSATION | 2 (4.7%) |  |  |
| COGNITIVE DISORDER | 1 (2.3%) |  |  |
| COMPLEX PARTIAL SEIZURES |  | 1 (2.4%) |  |
| COORDINATION ABNORMAL |  | 1 (2.4%) |  |
| DIZZINESS | 17 (39.5%) | 14 (33.3%) | 3 (18.8%) |
| HEADACHE | 9 (20.9%) | 4 (9.5%) | 8 (50%) |
| HEMIANOPIA HOMONYMOUS |  | 1 (2.4%) |  |
| HYPERSOMNIA | 1 (2.3%) |  |  |
| LETHARGY | 1 (2.3%) | 1 (2.4%) |  |
| PARAESTHESIA | 1 (2.3%) |  |  |
| PARAESTHESIA ORAL |  | 1 (2.4%) |  |
| PARKINSON'S DISEASE |  |  | 1 (6.2%) |
| PAROSMIA | 2 (4.7%) |  |  |
| PARTIAL SEIZURES WITH SECONDARY GENERALISATION | 1 (2.3%) |  |  |
| PSYCHOMOTOR HYPERACTIVITY |  |  | 1 (6.2%) |
| SOMNOLENCE | 1 (2.3%) | 5 (11.9%) | 3 (18.8%) |
| STUPOR |  | 1 (2.4%) |  |
| SYNCOPE | 3 (7%) | 7 (16.7%) |  |
| SYNCOPE VASOVAGAL | 1 (2.3%) |  |  |
| TRANSIENT ISCHAEMIC ATTACK | 1 (2.3%) | 3 (7.1%) |  |
| PSYCHIATRIC DISORDERS | 11 (100%) | 15 (100%) | 14 (100%) |
| AGITATION |  | 3 (20%) | 2 (14.3%) |
| ANXIETY |  | 4 (26.7%) | 2 (14.3%) |
| COMPLETED SUICIDE |  |  | 1 (7.1%) |
| CONFUSIONAL STATE | 1 (9.1%) | 3 (20%) | 2 (14.3%) |
| DELIRIUM | 1 (9.1%) |  |  |
| DELUSION | 1 (9.1%) |  | 1 (7.1%) |
| DEPRESSED MOOD | 1 (9.1%) | 2 (13.3%) |  |
| DISORIENTATION |  |  | 1 (7.1%) |
| HALLUCINATION | 1 (9.1%) |  |  |
| HALLUCINATION, VISUAL | 1 (9.1%) |  |  |
| INSOMNIA | 2 (18.2%) |  | 3 (21.4%) |
| IRRITABILITY |  | 1 (6.7%) | 2 (14.3%) |
| LIBIDO DECREASED | 1 (9.1%) |  |  |
| LISTLESS | 1 (9.1%) |  |  |
| NIGHTMARE | 1 (9.1%) |  |  |
| RESTLESSNESS |  | 2 (13.3%) |  |
| RENAL AND URINARY DISORDERS | 4 (100%) | 4 (100%) | 5 (100%) |
| CALCULUS URETHRAL | 1 (25%) |  |  |
| DYSURIA |  | 1 (25%) | 1 (20%) |
| ENURESIS |  | 1 (25%) |  |
| INCONTINENCE |  | 1 (25%) |  |
| MICTURITION URGENCY | 2 (50%) | 1 (25%) | 1 (20%) |
| NEPHROLITHIASIS | 1 (25%) |  | 1 (20%) |
| POLLAKIURIA |  |  | 2 (40%) |
| REPRODUCTIVE SYSTEM AND BREAST DISORDERS | 1 (100%) |  | 4 (100%) |
| BENIGN PROSTATIC HYPERPLASIA | 1 (100%) |  | 2 (50%) |
| PELVIC PAIN |  |  | 2 (50%) |
| RESPIRATORY, THORACIC AND MEDIASTINAL DISORDERS | 22 (100%) | 16 (100%) | 15 (100%) |
| ALLERGIC GRANULOMATOUS ANGIITIS | 1 (4.5%) |  |  |
| COUGH | 7 (31.8%) | 9 (56.2%) | 4 (26.7%) |
| DYSPHONIA |  | 1 (6.2%) |  |
| DYSPNOEA | 1 (4.5%) | 1 (6.2%) | 1 (6.7%) |
| EMPHYSEMA |  |  | 1 (6.7%) |
| EPISTAXIS | 2 (9.1%) | 1 (6.2%) |  |
| HAEMOPTYSIS |  |  | 2 (13.3%) |
| NASAL CONGESTION | 4 (18.2%) | 1 (6.2%) | 3 (20%) |
| PHARYNGEAL ERYTHEMA | 2 (9.1%) |  |  |
| PHARYNGOLARYNGEAL PAIN | 1 (4.5%) | 1 (6.2%) |  |
| POSTNASAL DRIP |  |  | 2 (13.3%) |
| PRODUCTIVE COUGH | 1 (4.5%) |  |  |
| RALES |  |  | 2 (13.3%) |
| RESPIRATORY TRACT CONGESTION | 1 (4.5%) |  |  |
| RHINORRHOEA | 2 (9.1%) | 2 (12.5%) |  |
| SKIN AND SUBCUTANEOUS TISSUE DISORDERS | 111 (100%) | 118 (100%) | 47 (100%) |
| ACTINIC KERATOSIS | 1 (0.9%) |  |  |
| ALOPECIA |  |  | 1 (2.1%) |
| BLISTER | 2 (1.8%) | 8 (6.8%) |  |
| COLD SWEAT |  |  | 3 (6.4%) |
| DERMATITIS ATOPIC |  |  | 1 (2.1%) |
| DERMATITIS CONTACT |  | 2 (1.7%) |  |
| DRUG ERUPTION |  |  | 1 (2.1%) |
| ERYTHEMA | 22 (19.8%) | 24 (20.3%) | 13 (27.7%) |
| HYPERHIDROSIS | 10 (9%) | 5 (4.2%) | 2 (4.3%) |
| PRURITUS | 38 (34.2%) | 35 (29.7%) | 11 (23.4%) |
| PRURITUS GENERALISED | 1 (0.9%) | 4 (3.4%) |  |
| RASH | 18 (16.2%) | 18 (15.3%) | 9 (19.1%) |
| RASH ERYTHEMATOUS |  | 2 (1.7%) |  |
| RASH MACULO-PAPULAR | 3 (2.7%) |  |  |
| RASH PAPULAR | 2 (1.8%) |  |  |
| RASH PRURITIC | 3 (2.7%) | 2 (1.7%) |  |
| SKIN EXFOLIATION |  | 2 (1.7%) |  |
| SKIN IRRITATION | 8 (7.2%) | 13 (11%) | 4 (8.5%) |
| SKIN ODOUR ABNORMAL | 1 (0.9%) |  |  |
| SKIN ULCER |  |  | 2 (4.3%) |
| URTICARIA | 2 (1.8%) | 3 (2.5%) |  |
| SOCIAL CIRCUMSTANCES | 1 (100%) |  |  |
| ALCOHOL USE | 1 (100%) |  |  |
| SURGICAL AND MEDICAL PROCEDURES | 2 (100%) | 1 (100%) | 2 (100%) |
| ACROCHORDON EXCISION | 1 (50%) |  |  |
| CATARACT OPERATION |  | 1 (100%) | 1 (50%) |
| EYE LASER SURGERY |  |  | 1 (50%) |
| SKIN LESION EXCISION | 1 (50%) |  |  |
| VASCULAR DISORDERS | 1 (100%) | 5 (100%) | 7 (100%) |
| HOT FLUSH |  | 1 (20%) |  |
| HYPERTENSION |  | 3 (60%) | 2 (28.6%) |
| HYPOTENSION |  | 1 (20%) | 3 (42.9%) |
| ORTHOSTATIC HYPOTENSION |  |  | 2 (28.6%) |
| WOUND HAEMORRHAGE | 1 (100%) |  |  |
| Abbreviations and comments | | | |
| Program name / Run date / Data cut date | | | |