

Table S1. Keywords selected from the particular “Reaction Groups” of EV database – part 1.

Ear and labyrinth disorders	Eye disorders	Gastrointestinal disorders	General disorders and administration site conditions	Immune system disorders
Ear swelling	Conjunctivitis allergic	Oral mucosa erosion*	Localised oedema ³	Allergic oedema ³
	Periorbital oedema ²	Oral mucosal blistering*	Infusion site rash	Allergic reaction to excipient
	Eye oedema ²	Oral mucosal exfoliation*	Injection site rash	Anaphylactic reaction**
	Eye swelling	Tongue blistering*	Oedema peripheral ³	Anaphylactic shock**
	Eyelid oedema ²	Stomatitis necrotising*	Peripheral swelling	Anaphylactoid reaction**
	Orbital oedema ²	Swollen tongue	Swelling	Anaphylactoid shock**
		Tongue exfoliation*	Face oedema ¹	Drug hypersensitivity
		Tongue ulceration*	Generalised oedema ³	Hypersensitivity
		Allergic colitis	Infusion site urticaria	Type I hypersensitivity
		Gastrointestinal oedema ³	Injection site swelling	Type IV hypersensitivity reaction
		Lip oedema ¹	Infusion site oedema ³	Type III immune complex mediated reaction
		Lip swelling	Oedema ³	
		Mouth swelling	Oedema mucosal ³	
		Oedema mouth ¹		
	Palatal oedema ¹			
	Palatal swelling			
	Tongue oedema ¹			

*keywords corresponding to prodromes or symptoms of Steven Johnson Syndrome/ Toxic Epidermal Necrolysis; ** keywords corresponding to diagnosis of anaphylaxis; ¹ keywords corresponding to head, neck or respiratory tract oedema excluding orbital or periorbital area; ² keywords corresponding to orbital or periorbital oedema; ³ keywords corresponding to oedema in other or unspecified localisation.

Table S2. Keywords selected from the particular “Reaction Groups” of EV database – part 2.

Investigations	Reproductive system and breast disorders	Respiratory, thoracic and mediastinal disorders	Vascular disorders
Eosinophil count increased	Genital erosion*	Catarrh	Bronchial obstruction
Eosinophil percentage increased	Genital rash	Rhinorrhoea	Bronchial oedema ¹
			Hot flush

Peak expiratory flow rate decreased	Genital swelling	Sinonasal obstruction	Bronchospasm	Shock
Allergy test positive	Genital ulceration	Sinus congestion	Cough	Shock symptom
	Oedema genital ³	Sneezing	Epiglottic oedema ¹	
	Perineal ulceration	Pharyngeal erosion	Laryngeal oedema ¹	
	Scrotal oedema ³	Status asthmaticus	Laryngeal stenosis	
	Scrotal swelling	Upper airway obstruction	Laryngospasm	
	Vulvar oedema ³	Upper respiratory tract congestion	Laryngotracheal oedema ¹	
	Vulvar ulceration*	Allergic cough	Nasal oedema ¹	
	Vulvovaginal swelling	Asthmatic crisis	Oropharyngeal swelling	
	Vulvar erosion*	Bronchial disorder	Pharyngeal oedema ¹	
		Bronchial hyperreactivity	Respiratory tract oedema ¹	
			Tracheal oedema ¹	
			Wheezing	

* keywords corresponding to prodromes or symptoms of Stevens-Johnson Syndrome/ Toxic Epidermal Necrolysis; ¹ keywords corresponding to head, neck or respiratory tract oedema excluding orbital or periorbital area; ² keywords corresponding to orbital or periorbital oedema; ³ keywords corresponding to oedema in other or unspecified localisation.

Table S3. Keywords selected from the particular “Reaction Groups” of EV database – part 3.

Skin and subcutaneous tissue disorders		
Mucosal erosion	Vasculitic rash	Rash
Mucosal exfoliation	Acute cutaneous lupus erythematosus	Rash erythematous
Mucosal ulceration	Acute generalised exanthematous pustulosis	Rash follicular
Blister rupture	Angioedema	Rash generalised
Circumoral oedema ³	Blister	Rash macular
Erythema multiforme	Dermatitis bullous	Rash maculo-papular
Fixed eruption	Dermatitis exfoliative	Rash morbilliform
Mucocutaneous rash	Dermatitis exfoliative generalised	Rash papular
Nikolsky's sign*	Drug eruption	Rash papulosquamous
Photosensitivity reaction	Drug reaction with eosinophilia and systemic symptoms	Rash pruritic
Skin erosion	Eczema vesicular	Rash rubelliform
Symmetrical drug-related intertriginous and flexural exanthema	Epidermal necrosis*	Rash scarlatiniform
Toxic epidermal necrolysis*	Erythema	Rash vesicular
Toxic skin eruption*	Exfoliative rash*	Skin exfoliation*

Urticaria	Generalised erythema	Skin oedema ³
Urticaria papular	Hypersensitivity vasculitis	Skin swelling
Urticaria vesiculosa	Swelling face	Stevens-Johnson Syndrome*

*keywords corresponding to prodromes or symptoms of Stevens-Johnson Syndrome/ Toxic Epidermal Necrolysis; ¹ keywords corresponding to head, neck or respiratory tract oedema excluding orbital or periorbital area; ² keywords corresponding to orbital or periorbital oedema; ³ keywords corresponding to oedema in other or unspecified localisation.

Table S4. The hypersensitivity symptoms reported as AE with frequency <1.1%.

Adverse event	No of reports	Adverse event	No of reports
Erythema multiforme	46	Skin erosion	5
Rash pruritic	45	Epidermal necrosis	5
Shock	44	Oral mucosal blistering	4
Bronchospasm	43	Circumoral oedema	4
Acute generalised exanthematous pustulosis	42	Rash vesicular	4
Laryngeal oedema	41	Genital ulceration	3
Wheezing	39	Allergic cough	3
Swelling	37	Laryngospasm	3
Lip oedema	36	Nasal oedema	3
Drug eruption	34	Mucosal ulceration	3
Oedema peripheral	32	Photosensitivity reaction	3
Tongue oedema	31	Allergic oedema	2
Pharyngeal oedema	31	Anaphylactoid shock Type IV	2
Flushing	31	hypersensitivity reaction	2
Dermatitis bullous	30	Eosinophil count increased	2
Oedema	29	Genital erosion	2
Rash macular	28	Sinus congestion	2
Swollen tongue	27	Bronchial hyperreactivity	2
Hypersensitivity vasculitis	26	Tracheal oedema	2
Anaphylactoid reaction	25	Symmetrical drug-related intertriginous and flexural exanthema	2
Drug reaction with eosinophilia and systemic symptoms	25	Rash scarlatiniform	2
Peripheral swelling	24	Skin swelling	2
Oedema mucosal	19	Oral mucosal exfoliation	1
Rash papular	19	Tongue blistering	1

Skin exfoliation	18	Stomatitis necrotising	1
Type I hypersensitivity	17	Tongue ulceration	1
Vulvar erosion	16	Palatal swelling	1
Rhinorrhoea	16	Injection site swelling	1
Generalised erythema	15	Allergic reaction to excipient	1
Localised oedema	14	Type III immune complex mediated reaction	1
Hot flush	14	Peak expiratory flow rate decreased	1
Shock symptom	12	Genital rash	1
Dermatitis exfoliative	11	Perineal ulceration	1
Toxic skin eruption	11	Scrotal oedema	1
Ear swelling	10	Sneezing	9
Palatal oedema	9	Scrotal swelling	1
Asthmatic crisis	5	Catarrh	1
Respiratory tract oedema	8	Upper respiratory tract congestion	1
Nikolsky's sign	8	Epiglottic oedema	1
Mouth swelling	7	Oropharyngeal swelling	1
Oedema mouth	7	Mucosal exfoliation	1
Rash morbilliform	6	Blister rupture	1
Eye oedema	5	Dermatitis exfoliative generalised	1
Oral mucosa erosion	5	Urticaria vesiculosa	1
Mucosal erosion	5		