

SUPPLEMENTARY APPENDIX

Efficacy and safety of 0.1% cyclosporine A cationic emulsion in dry eye disease: a pooled analysis of two double-masked, randomized, vehicle-controlled phase III clinical studies

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RESULTS

Supplementary Figure 1 SICCANOVE and SANSIKA study designs; EudraCT Numbers: 2007-000029-23 (SICCANOVE), 2011-000160-97 (SANSIKA). All patients who received study drug in the double-masked treatment (DMT) periods of the SICCANOVE and SANSIKA studies were included in the pooled analysis. Patients were allowed to use unpreserved artificial tears up to 6 times daily in the SICCANOVE study and as frequently as needed throughout the SANSIKA study. CFS scores based on modified Oxford scale. AT, artificial tears; CFS, corneal fluorescein staining; CsA CE, 0.1% (1 mg/mL) cyclosporine A in a cationic emulsion; DED, dry eye disease; DMT, double-masked treatment; FAS, full analysis set; HLA-DR, human leukocyte antigen-DR; OSDI, Ocular Surface Disease Index; QD, once daily. **[figure is presented separately]**

Supplementary Table 1 Drug-related TEAEs observed in the pooled patient population

	CsA CE n=396	Vehicle n=340
Drug-related TEAEs, n (%)	142 (35.9)	69 (20.3)
Severity of drug-related TEAEs, n (%)		
Mild	39 (9.8)	22 (6.5)
Moderate	17 (4.3)	12 (3.5)
Severe	86 (21.7)	35 (10.3)
Drug-related ocular TEAEs, n (%)	139 (35.1)	60 (17.6)
Serious drug-related ocular TEAEs, n (%)	1 (0.3)	1 (0.3)
Severe drug-related ocular TEAEs, n (%)	86 (21.7)	34 (10)
Study discontinuation due to drug-related ocular TEAEs, n (%)	37 (9.3)	20 (5.9)
Most common drug-related TEAEs (occurring in $\geq 2\%$ of patients in either group), n (%)		
Instillation site pain	48 (12.1)	9 (2.6)
Eye irritation	40 (10.1)	9 (2.6)
Instillation site irritation	20 (5.1)	4 (1.2)
Eye pain	16 (4.0)	9 (2.6)
Erythema of eyelid	9 (2.3)	5 (1.5)
Lacrimation increased	9 (2.3)	1 (0.3)
Visual acuity reduced	5 (1.3)	7 (2.1)

Conjunctival hyperemia	8 (2.0)	3 (0.9)
Instillation site erythema	8 (2.0)	0 (0.0)

CsA CE, 0.1% (1 mg/mL) cyclosporine A in a cationic emulsion; TEAEs, treatment-emergent adverse events.

INVESTIGATORS IN THE SICCANOVE AND SANSIKA STUDIES			
Country	Investigator	SICCANOVE	SANSIKA
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UK	Dr. Balasubramaniam Ilango	X	
UK	Dr. Mohan V. Mundasad	X	
UK	Dr. Guy T. Smith	X	