## IN THE UNITED STATES DISTRICT COURT FOR THE DISTRICT OF DELAWARE

MEDA PHARMACEUTICALS INC. and CIPLA LTD.,	)	
Plaintiffs,	) )	C A N. 14 01452 I DO
v.	)	C.A. No. 14-01453-LPS
APOTEX INC. and APOTEX CORP.,	)	
Defendants.	)	
	)	

### PLAINTIFFS' POST-TRIAL SUR-REPLY BRIEF ON OBJECTIVE INDICIA OF NONOBVIOUSNESS

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	A.	UNEXPECTED RESULTS: A POSA in 2002 would not have expected Dymista®'s clinical benefits and successful formulation.	5
		1. Dymista®'s increased efficacy, accelerated onset, and reduced side effects were not expected in 2002.	5
		2. Dymista <sup>®</sup> 's benefits are differences in "kind."	8
		3. A POSA would not have expected that a single formulation like Dymista <sup>®</sup> could be made suitable for nasal administration	10
	В.	FAILURE OF OTHERS: At least three sophisticated pharmaceutical companies failed to develop an antihistamine/steroid fixed-dose formulation.	11
	C.	ACQUIESCENCE: Meda took a license from Cipla for the Patents-in-Suit to market Dymista <sup>®</sup> .	15
	D.	SKEPTICISM: Both Meda and FDA doubted whether a fixed-dose antihistamine/steroid combination would work	15
		Clinical skepticism, prior to and well after the invention date, confirms that a POSA would not have anticipated success.	16
		2. Industry participants were skeptical that an antihistamine/steroid combination could be formulated	18
	E.	UNMET NEED: Dymista® satisfied a long-felt but previously unmet need for better, faster, and safer treatment of AR	19
	F.	COMMERCIAL SUCCESS: Apotex has failed to overcome evidence of Dymista®'s commercial success	19
	G.	COPYING: Duonase (the version of Dymista <sup>®</sup> in India) was immediately and widely copied by multiple pharmaceutical companies.	21
	H.	PRAISE: Dymista <sup>®</sup> has been received by the medical community as best treatment for AR available.	22



II.		RE WERE NO "BLOCKING PATENTS" TO NEGATE ANY OBJECTIVE CIA OF NONOBVIOUSNESS	23
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