Efficiency and Safety of Ripasudil Hydrochloride Hydrate for Uveitis-associated Ocular Hypertension

Kei Takayama¹¶, Mami Yasuda, ¹¶Takayuki Kanda¹*, Manzo Taguchi¹, Hideaki Someya¹, Toshiaki Murata¹, and Masaru Takeuchi¹ ¹Department of Ophthalmology, National Defense Medical College, Tokorozawa, Japan ^{Financial interests: none}

Background:

Ripasudil hydrochloride hydrate (K-115) is an eye drop developed for the treatment of glaucoma and ocular hypertension in Japan. The aim of this study to evaluate the efficiency and safety of ripasudil hydrochloride hydrate for uveitis-associated ocular hypertension (OHT) and compare the outcomes of different uveitis.

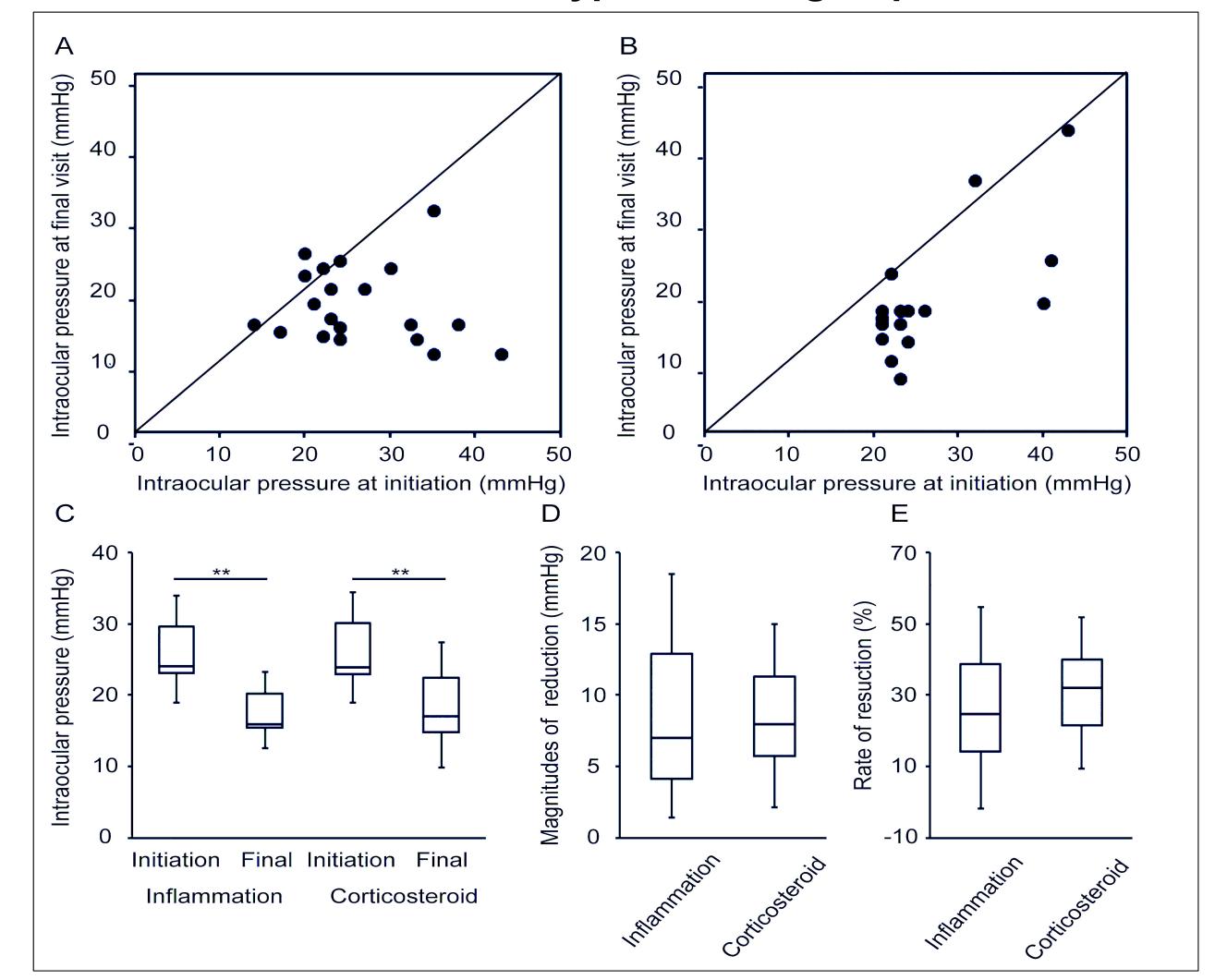
Patients & Methods: Clinical records of 36 eyes of 27 patients with uveitis-associated OHT (18 eyes with granulomatous uveitis and 10 eyes with non-granulomatous uveitis) who were treated by K-115 combined with or without other antiglaucoma agents were retrospectively reviewed. Intraocular pressure (IOP), aqueous flare, and posterior inflammation scores were compared before and after treatment with K-115.

<u>Results:</u>

Table 1. Patients' characteristics in the inflammation- and corticosteroidinduced ocular hypertension groups

Inflammation Corticosteroi P

Figure 1. Changes in intraocular pressure in the inflammation- and corticosteroid-induced ocular hypertension groups



Number of eyes (people)	20 (16)	16 (11)	
Male/female (people)	5/11	6/5	
Mean age (years)	62.1 ± 13.2	63.0 ± 15.2	0.36*
Mean observation duration (months)	5.2 ± 3.0	5.2 ± 2.7	0.43*
Mean BCVA (logMAR)	0.47 ± 0.82	0.48 ± 0.80	0.44*
Diagnosis of uveitis (eyes)			
Scleritis-associated uveitis	5	2	
Behçet's disease		6	
Herpes uveitis	3		
Sarcoidosis	2	1	
Vogt–Koyanagi–Harada disease		2	
Cytomegalovirus-associated uveitis	1		
Unknown	9	5	
Prescribed anti-glaucoma agents			
Prostaglandin analogues	15	13	
β-blockers	12	9	0.71#
Carbonic anhydrase inhibitors	7	5	U.7 I"
α2-agonists	5	8	

Figure 2. Changes in anterior aqueous flare in the inflammation- and corticosteroid-induced ocular hypertension groups

(A) IOP at the initiation and final visit in the inflammation group. (B) IOP at the initiation and final visit in the corticosteroid group. (C) The mean IOP at the initiation and final visit in the two groups. The mean IOP was reduced significantly by K-115 initiation in both groups. (D) The magnitudes of IOP reduction in the two groups. No significant differences were found between the two groups (E) The rates of IOP reduction in the two groups. *: P < 0.01

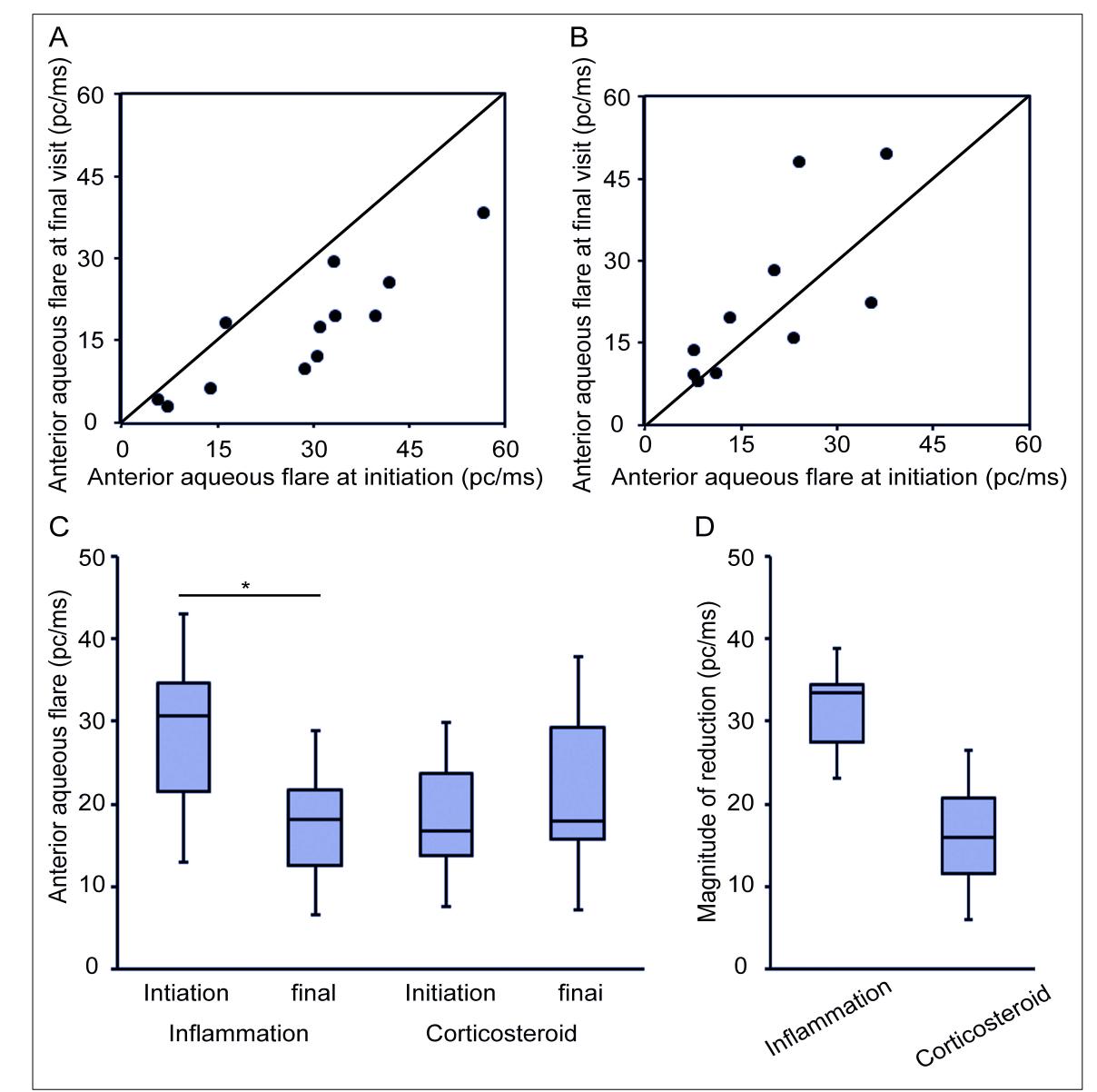


Figure 3. Correlations between intraocular pressure and anterior aqueous flare in the inflammation- and corticosteroid-induced ocular hypertension groups

(A) There was a tendency of correlation (pc/ms) (pc/ms) 50 between IOP reduction reduction reduction and anterior flare 25 35 decrease in the inflammation group. flare flare 20 (B) There was no aqueous aqueous significant correlation 5 between IOP reduction P = 0.078 and anterior flare Anterior = 0.44-20 10 change in the -20 40 -10 20 50 15 35 10 IOP reduction (mmHg) corticosteroid group. IOP reduction (mmHg) Table 2 Factors affecting changes in IOP and aqueous flare

Table 2 Factors anecting changes	in IOP and aq	ueous nare			
	ΙΟ	IOP		Flare	
	Р	β	Р	β	
Inflammation/corticosteroid	0.95		0.24		
Age	0.51		0.24		
Sex	0.57		0.45		
Granulomatous/non- granulomatous	0.93		0.86		
Prescribed anti-glaucoma agents					
Prostaglandin analogues	0.23		0.65		
β-blockers	0.68		0.21		
Carbon anhydrase inhibitors	0.94		0.63		
α2-agonists	0.64		0.03	-18.30	
Aqueous flare at the initiation	0.45		0.21		
IOP at the initiation	<0.001	1.40	0.02	0.56	

(A) Anterior aqueous flare at the initiation and final visit in the inflammation group. (B) Anterior aqueous flare at the initiation and final visit in the corticosteroid group. (C) The mean anterior aqueous flare in the two groups at the initiation and final visit. In the inflammation group, the mean anterior aqueous flare was significantly decreased. (D) The magnitude of change in anterior aqueous flare in the two groups. *: P< 0.05

Comments:

K-115 is effective in lowering IOP and is safe and well-tolerated in eyes with uveitis-associated OHT, including both inflammation- and corticosteroid-induced OHT. K-115 may suppress the anterior inflammation in eyes with inflammation-induced OHT and may have stronger effects in eyes with corticosteroid-induced OHT.

Conclusions:

K-115 decreased IOP in both inflammation- and corticosteroid-induced OHT associated with uveitis and played a synergistic role in reducing ocular inflammation in uveitis treatment.