

A close-up photograph of a human eye with a striking, vibrant blue iris. The eye is looking slightly to the right. The surrounding skin is fair and the eyelashes are dark and well-defined. The background is dark and out of focus.

ALTISSIMO Full-Data Analysis 12-Month Treatment Phase

Conference Call

May 12, 2021

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Forward-Looking Statements

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Today's Presenters



Fred Guerard
PharmD, CEO

- ✓ **Novartis**
Worldwide Head
Ophthalmology
- ✓ **Alcon**
Global Franchise Head
Pharmaceuticals
- ✓ Led extension of Novartis
ophthalmology pipeline:
Encore Vision, Lubricin®,
Luxturna®, Xiidra®



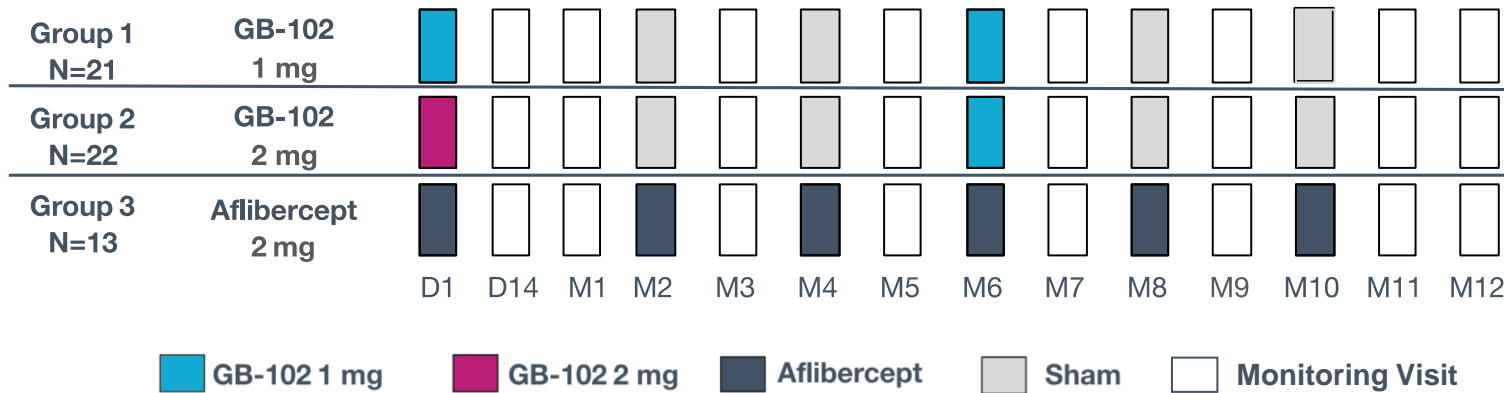
Parisa Zamiri
MD, PhD, CMO

- ✓ **Novartis**
VP and Global Head, Clinical
Development and
Therapeutic Area Head,
Ophthalmology
- ✓ Ophthalmologist from
Moorfield's Eye Hospital, UK
- ✓ Ocular immunologist from
Schepens Eye Research
Institute, a Harvard affiliated
institute

GB-102 Phase 2b trial in wet AMD (ALTISSIMO)



Primary endpoint: time to first use of additional supportive therapy



Population

- nAMD ≤ 18 months
- ≥ 3 prior anti-VEGF injections
- Response to treatment
- Anti-VEGF ≤ 21 days
- 20/20 - 20/200

Endpoints

- Time to first rescue
- BCVA
- CST (OCT)
- Adverse events

Extension study to provide information on GB-102 1mg beyond month 12

*6 patients withdrew for reasons unrelated to their treatment

Overall, GB-102 1 mg was safe and well tolerated

- ✓ **No drug-related serious adverse events**
- ✓ No Treatment Emergent Adverse Events (study eye or non-ocular) leading to drug discontinuation
- ✓ No adverse event requiring surgical intervention
- ✓ **No vision-threatening inflammation or increase in intraocular pressure**
- ✓ Majority of drug-related adverse events were mild to moderate
- ✓ Medication was detected in the anterior chamber (AC) in 7.8% of GB-102 1 mg injections, **a 79% reduction from ADAGIO**

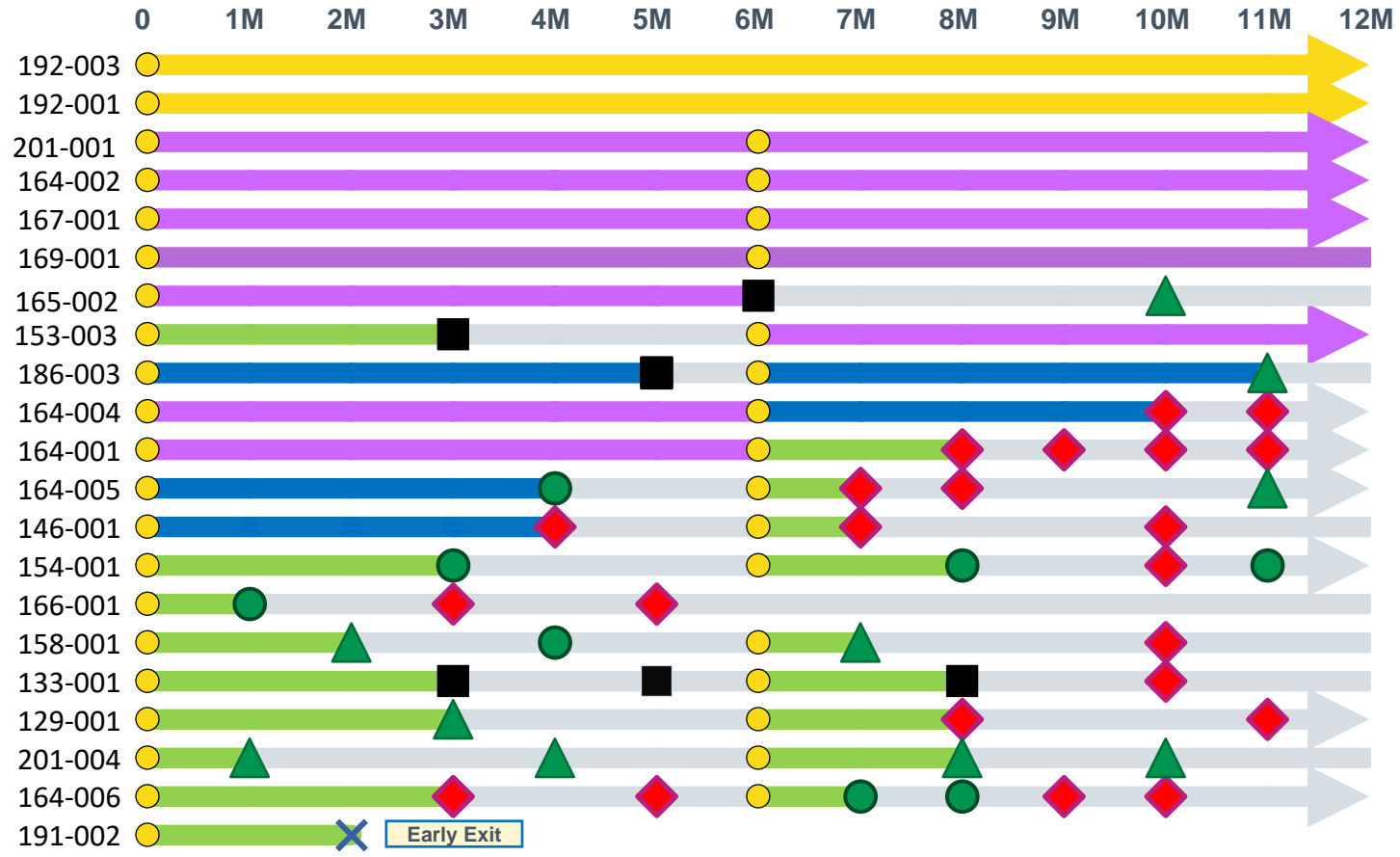
Optimization of manufacturing process led to 79% reduction in frequency of presence of medication in anterior chamber compared to ADAGIO

Drug-related TEAE (Study Eye)	aflibercept (N = 13)	GB-102 1mg/1mg (N = 21) [BL-6/6-12]*
Product residue present in AC	0	3 [2/1]
Vitreous floaters	0**	5 [5/1]
Product residue in the vitreous	0	1 [1/0]
Iritis	0	4 [2/3]
Uveitis	0	1 [0/1]
Vitritis	0	1 [0/1]
Vision blurred	0	1 [0/1]
Visual acuity reduced	0	2 [0/2]
Visual impairment	0	1 [1/0]
Retinal haemorrhage	0	1 [1/0]

*Some patients experienced the same AE during both 6 months period

**There were 2 cases of floaters in aflibercept group that were deemed not related

Median Time to Additional Supportive Therapy for GB-102 1 mg: 5 months



Time to additional supportive therapy						
GB-102 1mg	1M	2M	≥3M	≥4M	≥5M	≥6M
Dose 1	10%	10%	81%	57%	48%	43%
Dose 2	25%	31%	44%	44%	38%	31%

Duration:

- After 1st rescue
- ≤ 3M
- 4-5 M
- 6-7 M
- ≥ 8M

Dosing:

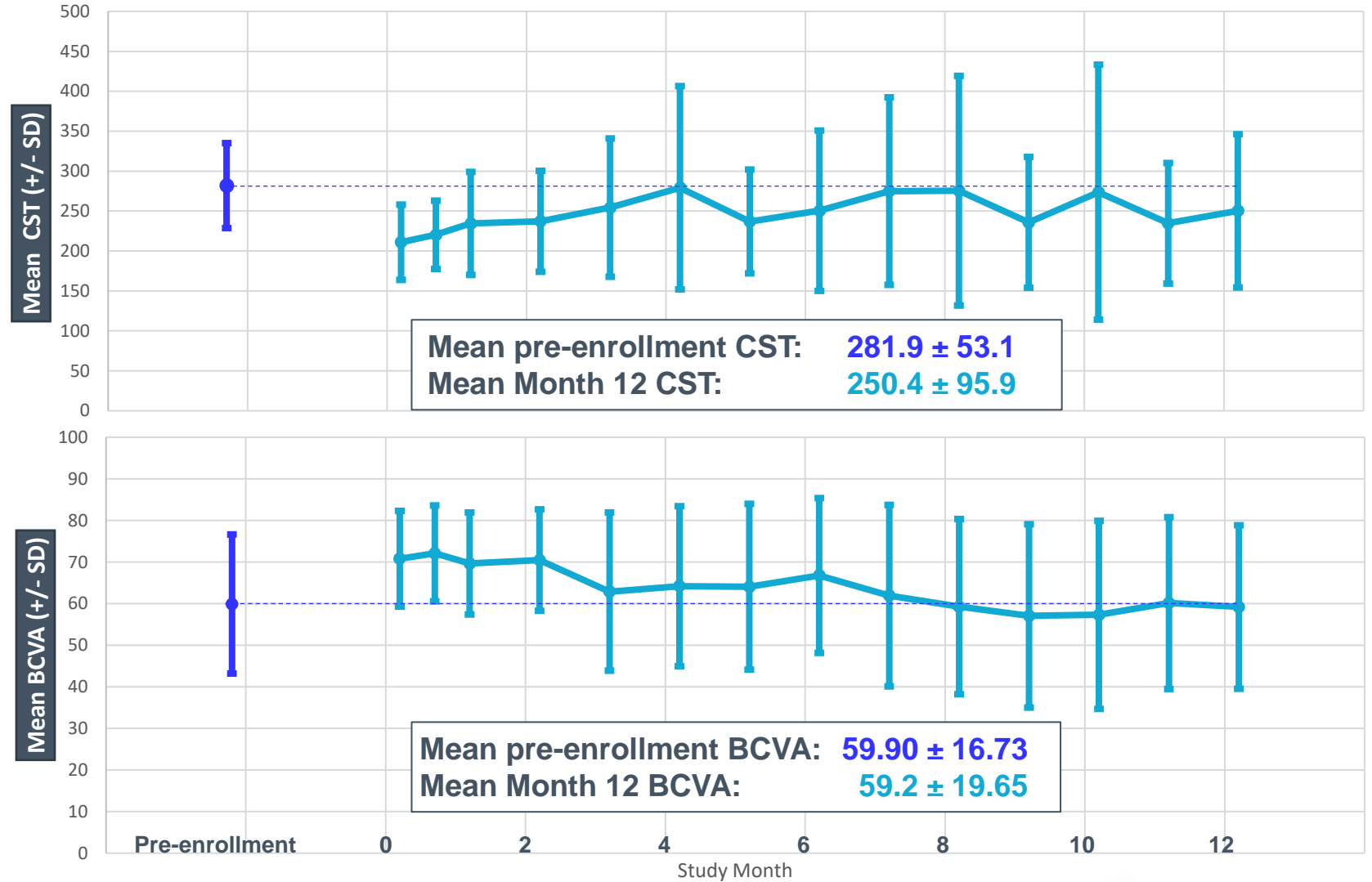
- Scheduled dosing
- Rescued solely due to BCVA
- Rescued solely due to CST
- Rescued due to BCVA and CST
- Rescued, but no criteria met

Treatment burden reduced by 58% while CST and BCVA was maintained

Injections per Year (GB-102 Arm)

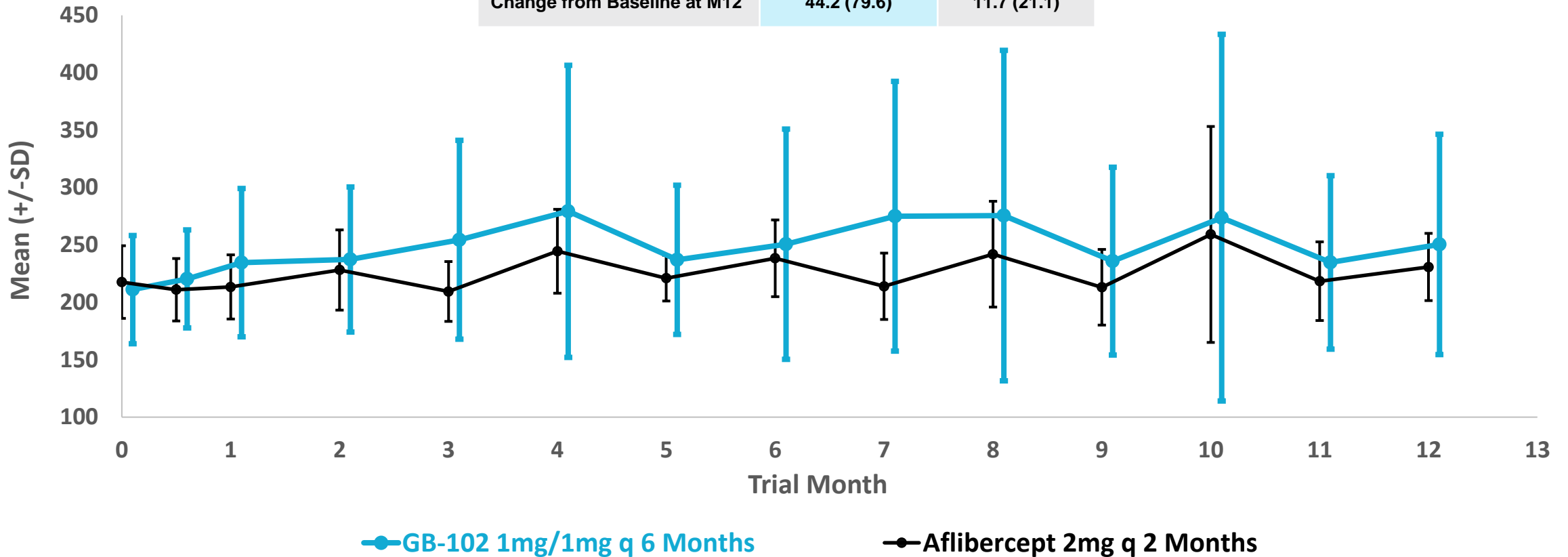
Mean prior anti-VEGF: 10.2

Mean on-trial anti-VEGF: 4.2



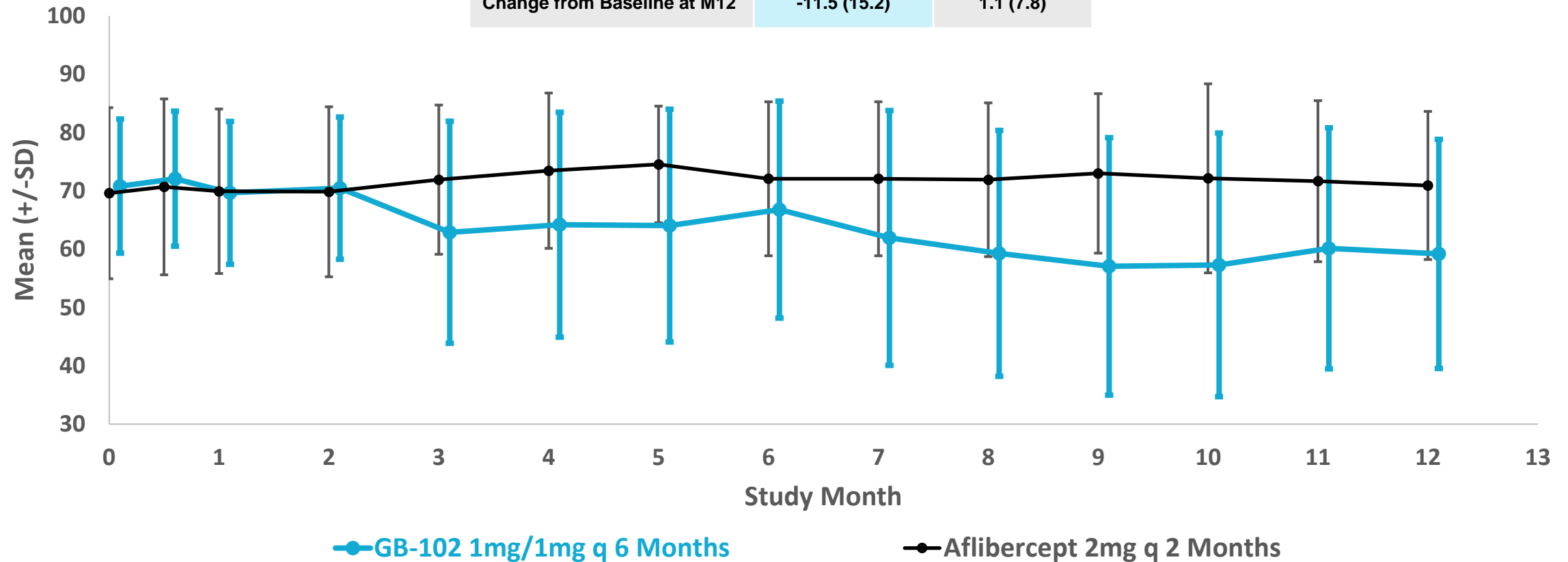
Control of CST with GB-102 1 mg given every 6 months was similar to that of bi-monthly aflibercept

Time Frame	GB-102 1mg/1mg Mean (SD)	aflibercept Mean (SD)
Change from Baseline at M6	44.3 (82.2)	19.3 (22.5)
Change from Baseline at M12	44.2 (79.6)	11.7 (21.1)



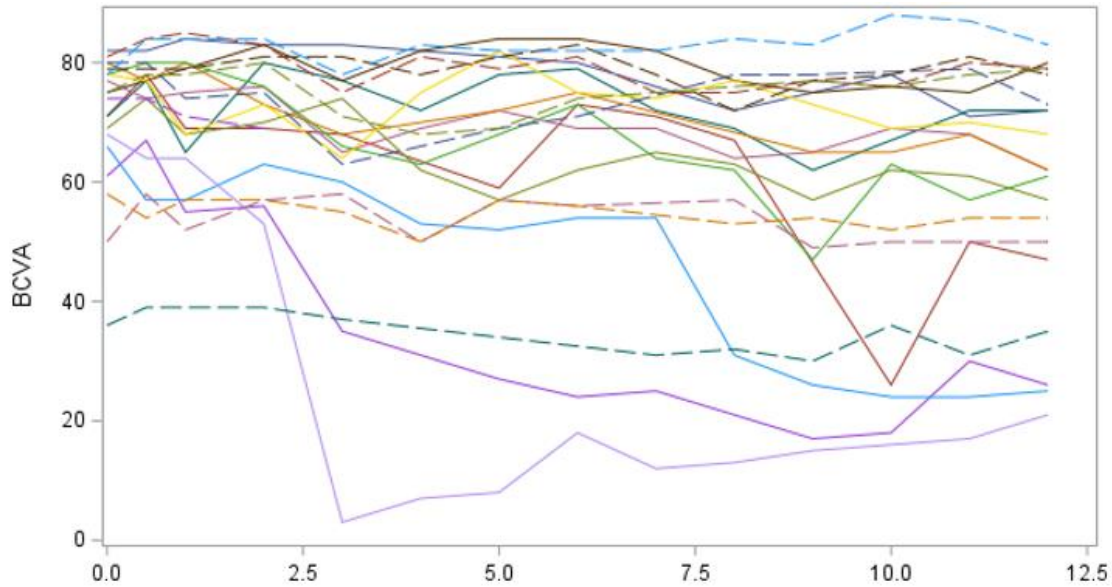
BCVA trended lower in GB-102 1 mg given every 6 months as compared with bi-monthly aflibercept — high standard deviation driven by 6 patients

Time Frame	GB-102 1mg/1mg Mean (SD)	aflibercept Mean (SD)
Change from Baseline at M6	-5.7 (14.7)	2.3 (5.1)
Change from Baseline at M12	-11.5 (15.2)	1.1 (7.8)



Opportunity to optimize clinical trial design and using enhanced formulation to deliver similar BCVA results to aflibercept

6 patients adversely impacted mean BCVA



Mean BCVA Letter Changes

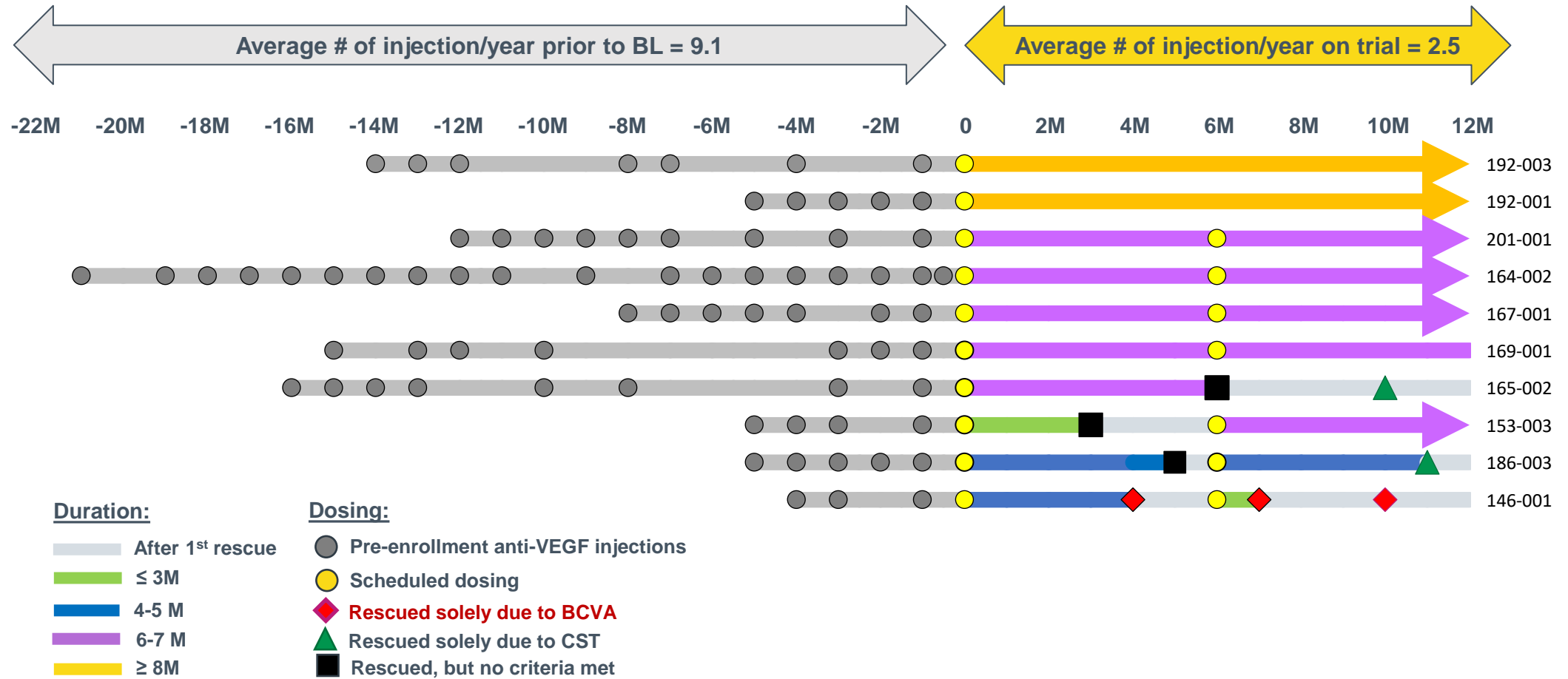
	@Baseline	M1-12	@M12*
Aflibercept	0.0	1.8	1.1
All GB-102 1mg	0.0	(7.4)	(11.5)
Subgroup of 6	0.0	(18.8)	(24.2)
Subgroup of 14	0.0	(2.7)	(6.0)
All GB-102 vs. aflibercept	0.0	(9.3)	(12.5)
Best 14 vs. aflibercept	0.0	(4.5)	(7.1)

Median BCVA Letter Changes

	@Baseline	M1-12	@M12
Aflibercept	0.0	1.0	1.0
All GB-102 1mg	0.0	(5.5)	(8.5)
Subgroup of 6	0.0	(12.5)	(21.0)
Subgroup of 14	0.0	(1.0)	(1.5)
All GB-102 vs. aflibercept	0.0	(6.5)	(9.5)
Best 14 vs. aflibercept	0.0	(2.0)	(2.5)

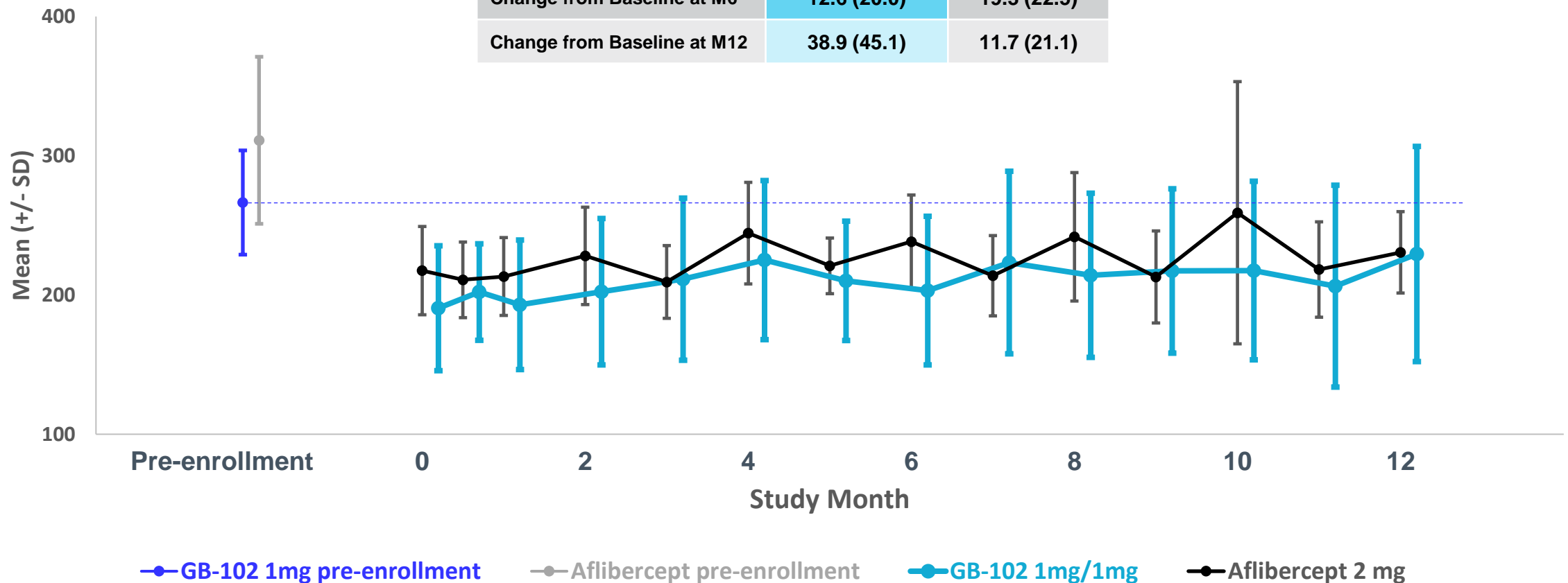
*M1-12 is the median of the medians BCVA loss across those visits

GB-102 1 mg reduced injection frequency of the top 10 patients by 72%



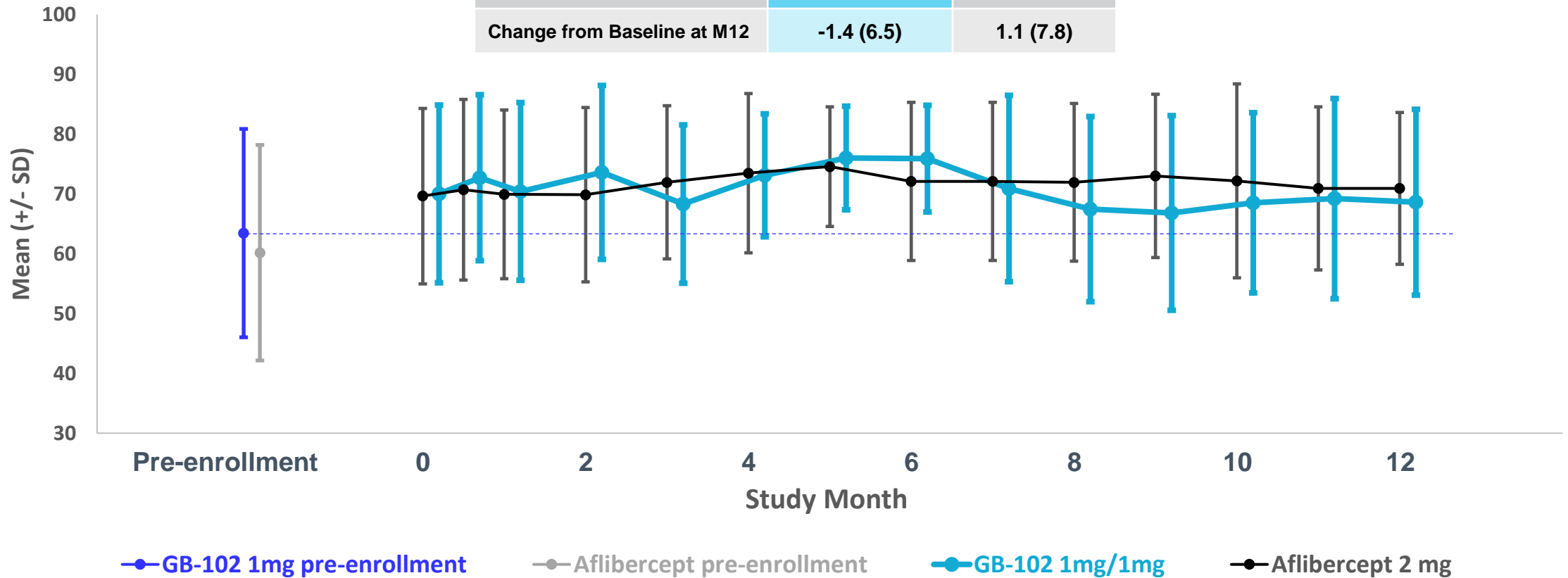
GB-102 1 mg controlled CST of top 10 patients whilst reducing magnitude of fluctuations over 12 months by 33%

Time Frame	GB-102 1mg/1mg Mean (SD)	aflibercept Mean (SD)
Change from Baseline at M6	12.6 (20.0)	19.3 (22.5)
Change from Baseline at M12	38.9 (45.1)	11.7 (21.1)



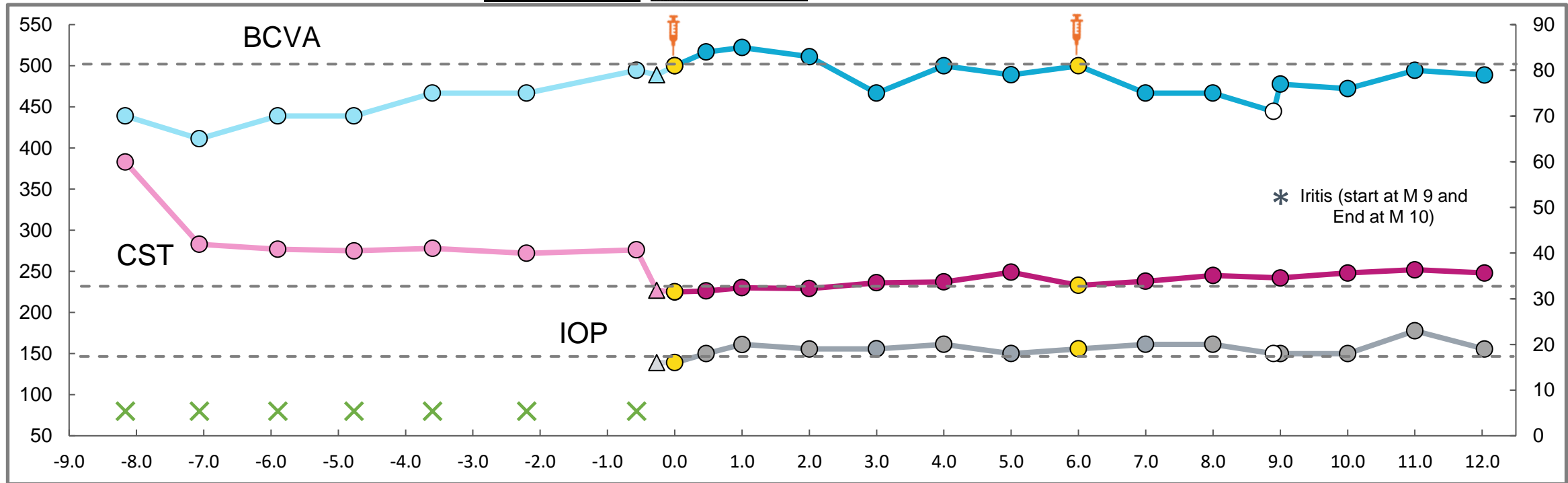
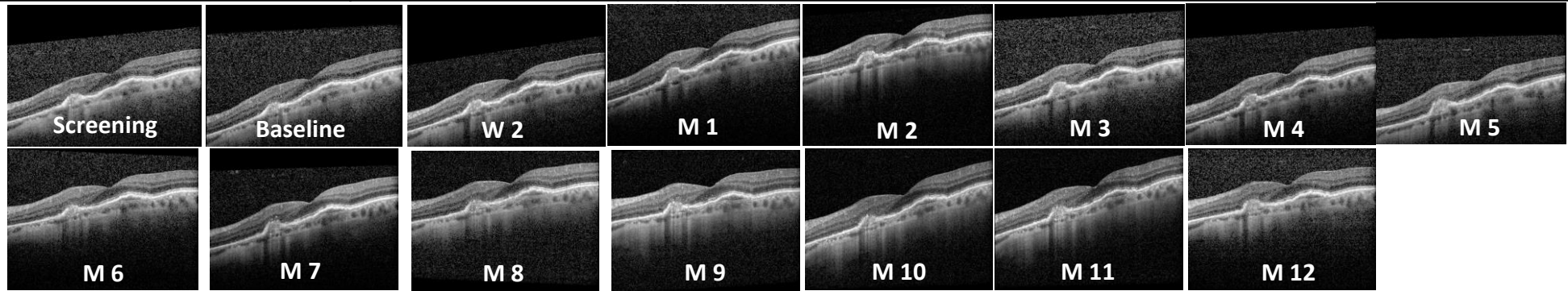
GB-102 1mg provided top 10 patients with BCVA control within non-inferiority margin* to aflibercept

Time Frame	GB-102 1mg/1mg Mean (SD)	aflibercept Mean (SD)
Change from Baseline at M6	2.1 (5.6)	2.3 (5.1)
Change from Baseline at M12	-1.4 (6.5)	1.1 (7.8)



*ALTISSIMO not powered to demonstrate non-inferiority

Subject ID: XXX-XXX	Demographic: 66/F/W	Lesion type: Minimally classic, CNV
Treatment/Study Eye: 1.0 mg GB-102/OS	Prior Injections: 7	Relevant Medical History: nAMD (OS), Cataract (OU), Dry AMD (OU), PVD (OU)



Actionable Findings from ALTISSIMO

Capitalize on good anatomical control and increase probability of success

Variable	ALTISSIMO	Action
Duration	48% of patients reached 6 Months	Refining criteria could reduce rescues by >50%
	Two hard-to-treat patients	Tighten inclusion criteria
Reduction in BCVA primarily driven by subgroup of six patients	Two treatment-unrelated AEs	Larger study will distribute these patients evenly
	Two events of particle dispersion	Optimized formulations should improve behavior
Reconstitution / Injection Procedure	Inconsistent aggregation	Faster aggregating microparticles or implants will reduce injection technique variability

ALTISSIMO Summary

- First IVT treatment to demonstrate 6-month duration for half of patients in controlled trial
- Safety and tolerability of GB-102 1 mg established
- Efficacy of sunitinib demonstrated by anatomical control similar to Eylea
- Lower trend in visual outcome is understood and can be addressed
- Work on optimized formulations of GB-102 initiated in 2020
- Prioritization of capital towards projects with potential for near-term catalysts

Thank You

