Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eAppendix. STROBE Reporting Guidelines: Additional Details and Exceptions

This manuscript followed the STROBE Reporting Guidelines for a cohort study, with additional details and a few exceptions described below.

- 1. **Setting:** Dates, including periods of recruitment and data collection, are not included due to attributability concerns for astronaut and cosmonaut crewmembers.
- 2. **Study size:** A sample size calculation was performed for the larger Fluid Shifts study and determined that a minimum sample size of 10 crewmembers was required. Fourteen crewmembers were included to account for any missing data.
- 3. **Descriptive Data:** Missing data are described below:
 - a. One subject dropped out of the study after flight day (FD) 50 for reasons unrelated to the study
 - b. Lower body negative pressure (LBNP) data was not included for two subjects:
 - i. One subject did not participate in inflight LBNP due to development of a retinal hemorrhage
 - ii. One subject had scan quality issues on FD50 and scheduling constraints on FD150
 - c. Five Russian subjects were unable to return for imaging 10 days after return to Earth (R+10) due to travel constraints

All other items on the STROBE checklist are addressed in the main text of the manuscript.

eTable 1. P Values for Change in Each Parameter During and After Spaceflight Relative to the Preflight Seated Value

	FD50 vs. Preflight	FD150 vs. Preflight	R+10 vs. Preflight	R+30 vs. Preflight	R+180 vs. Preflight
MRW (μm)	< .001	< .001	< .001	.007	.46
Cup volume (mm³)	< .001	< .001	< .001	< .001	> .99
BMO height (μm)	.06	.009	.78	.97	.24
TRT250 (μm)	< .001	< .001	< .001	.002	.36
TRT500 (μm)	< .001	< .001	< .001	.002	.24
TRT1000 (μm)	.72	.02	.27	.13	.02
TRT1500 (μm)	.33	.77	.46	.96	.007
MT500 (μm)	< .001	< .001	< .001	.003	.07
MT1500 (μm)	< .001	< .001	< .001	.38	.13
MT2500 (μm)	.11	.10	.06	.53	.41

MRW, minimum rim width; BMO, Bruch's membrane opening; TRT, total retinal thickness; TRT250, TRT from BMO to 250 μ m; TRT500, TRT from 250 to 500 μ m; TRT1000, TRT from 500 to 1000 μ m; TRT1500, TRT from 1000 to 1500 μ m; MT, macular thickness; MT500, MT from the fovea to 500 μ m; MT1500, MT from 500 to 1500 μ m; MT2500, MT from 1500 to 2500 μ m; FD, flight day; R+, days after return to Earth.

The R+10 time point includes only 8 subjects, as several international crewmembers did not return directly to Houston

eTable 2. Estimated Marginal Mean (95% CI) for Each Parameter Without and With Lower-Body Negative Pressure During Spaceflight

	FD50	FD50 + LBNP	FD150	FD150 + LBNP
MRW (μm)	386.4	385.6	398.6	402.1
	(349.2 – 423.5)	(348.3 – 422.8)	(361.4 – 435.8)	(364.9 – 439.3)
Cup volume (mm³)	0.167	0.168	0.158	0.155
	(0.076 – 0.258)	(0.077 – 0.259)	(0.067 – 0.249)	(0.063 – 0.246)
BMO height (μm)	-124.7	-125.0	-127.2	-122.2
	(-156.3 – -93.0)	(-156.7 – -93.3)	(-158.8 – -95.6)	(-153.9 – -90.6)
TRT250 (μm)	413.0	412.9	424.4	427.3
	(387.5 – 438.6)	(387.3 – 438.5)	(398.9 – 450.0)	(401.7 – 452.9)
TRT500 (μm)	381.7	381.0	387.6	388.6
	(365.4 – 398.1)	(364.6 – 397.4)	(371.2 – 404.0)	(372.2 – 405.0)
TRT1000 (μm)	339.9	339.6	341.5	341.9
	(329.3 – 350.5)	(329.0 – 350.2)	(330.9 – 352.0)	(331.3 – 352.5)
TRT1500 (μm)	303.2	303.2	302.7	303.4
	(294.9 – 311.4)	(294.9 – 311.4)	(294.5 – 311.0)	(295.1 – 311.7)
MT500 (μm)	268.7	269.5	267.2	267.0
	(258.6 – 278.7)	(259.4 – 279.6)	(257.1 – 277.2)	(256.9 – 277.1)
MT1500 (μm)	345.5	345.9	344.4	344.2
	(336.6 – 354.4)	(336.9 – 354.8)	(335.5 – 353.4)	(335.3 – 353.2)
MT2500 (μm)	314.8	315.3	314.7	314.0
	(307.1 – 322.4)	(307.6 – 322.9)	(307.1 – 322.4)	(306.3 – 321.7)

MRW, minimum rim width; BMO, Bruch's membrane opening; TRT, total retinal thickness; TRT250, TRT from BMO to 250 μ m; TRT500, TRT from 250 to 500 μ m; TRT1000, TRT from 500 to 1000 μ m; TRT1500, TRT from 1000 to 1500 μ m; MT, macular thickness; MT500, MT from the fovea to 500 μ m; MT1500, MT from 500 to 1500 μ m; MT2500, MT from 1500 to 2500 μ m; FD, flight day; LBNP, lower body negative pressure.

eTable 3. P Values for Change in Each Parameter With LBNP and Between-Flight Days Without LBNP

	FD50 vs.	FD150 vs.	FD50 vs.
	FD50 + LBNP	FD150 + LBNP	FD150
MRW (μm)	.76	.20	< .001
Cup volume (mm³)	.85	.06	< .001
BMO height (μm)	.90	.07	.31
TRT250 (μm)	.93	.17	< .001
TRT500 (μm)	.63	.28	< .001
TRT1000 (μm)	.63	.43	.04
TRT1500 (μm)	.94	.15	.43
MT500 (μm)	.30	.84	.046
MT1500 (μm)	.54	.76	.06
MT2500 (μm)	.32	.18	.99

MRW, minimum rim width; BMO, Bruch's membrane opening; TRT, total retinal thickness; TRT250, TRT from BMO to 250 μ m; TRT500, TRT from 250 to 500 μ m; TRT1000, TRT from 500 to 1000 μ m; TRT1500, TRT from 1000 to 1500 μ m; MT, macular thickness; MT500, MT from the fovea to 500 μ m; MT1500, MT from 500 to 1500 μ m; MT2500, MT from 1500 to 2500 μ m; FD, flight day; LBNP, lower body negative pressure.

eTable 4. Estimated Marginal Mean (95% CI) for Each Parameter With Posture Changes Before and After Spaceflight

	Preflight Seated	Preflight Supine	Preflight HDT	R+10 Seated	R+10 Supine	R+10 HDT
MRW (μm)	359.5	360.8	363.6	380.7	380.2	383.5
	(330.3 – 388.6)	(331.6 – 389.9)	(334.4 – 392.7)	(351.4 – 410.0)	(351.0 – 409.5)	(354.2 – 412.7)
Cup volume (mm³)	0.193	0.197	0.190	0.162	0.163	0.160
	(0.102 – 0.284)	(0.106 – 0.288)	(0.099 – 0.281)	(0.071 – 0.253)	(0.072 – 0.255)	(0.069 – 0.251)
BMO height (μm)	-118.3	-122.7	-121.2	-114.0	-115.9	-112.9
	(-146.7 – -89.8)	(-151.2 – -94.3)	(-149.7 – -92.8)	(-142.6 – -85.3)	(-144.5 – -87.3)	(-141.5 – -84.2)
TRT250 (μm)	394.3	391.8	394.2	410.0	409.8	412.6
	(375.9 – 412.7)	<i>(373.4 – 410.1)</i>	(375.8 – 412.6)	(391.5 – 428.5)	(391.3 – 428.3)	(394.2 – 431.1)
TRT500 (μm)	370.9	370.2	370.9	381.1	379.9	381.2
	(358.6 – 383.2)	(357.9 – 382.5)	(358.6 – 383.2)	(368.7 – 393.5)	(367.5 – 392.3)	(368.8 – 393.5)
TRT1000 (μm)	337.6	337.1	336.9	340.5	340.2	339.5
	(327.7 – 347.5)	(327.2 – 347.0)	(327.0 – 346.8)	(330.6 – 350.4)	(330.3 – 350.1)	(329.6 – 349.4)
TRT1500 (μm)	302.5	302.1	301.7	302.4	302.4	302.3
	(294.2 – 310.8)	(293.8 – 310.5)	(293.4 – 310.1)	(294.1 – 310.8)	(294.0 – 310.8)	(294.0 – 310.7)
MT500 (μm)	272.4	272.0	271.1	267.6	266.7	265.9
	(262.7 – 282.1)	(262.3 – 281.7)	(261.4 – 280.9)	(257.8 – 277.3)	(257.0 – 276.5)	(256.1 – 275.6)
MT1500 (μm)	348.1	348.5	348.1	345.1	344.3	343.7
	(339.3 – 356.9)	(339.7 – 357.3)	(339.3 – 356.9)	(336.3 – 353.9)	(335.5 – 353.1)	(334.9 – 352.6)
MT2500 (μm)	315.7	316.3	316.0	314.7	314.3	313.6
	(307.9 – 323.5)	(308.5 – 324.2)	(308.2 – 323.8)	(306.8 – 322.5)	(306.5 – 322.2)	(305.8 – 321.4)

MRW, minimum rim width; BMO, Bruch's membrane opening; TRT, total retinal thickness; TRT250, TRT from BMO to 250 μ m; TRT500, TRT from 250 to 500 μ m; TRT1000, TRT from 500 to 1000 μ m; TRT1500, TRT from 1000 to 1500 μ m; MT, macular thickness; MT500, MT from the fovea to 500 μ m; MT1500, MT from 500 to 1500 μ m; MT2500, MT from 1500 to 2500 μ m; HDT, head-down tilt; R+, days after return to Earth.

The R+10 time point includes only 8 subjects, as several international crewmembers did not return directly to Houston.