# 1Self-corrected Hemodynamic Analysis of Blood Viscosity for Detection2of Endothelial Injury in Atherosclerosis using MRI

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#### **Supplementary Materials**

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### 4 Section 1: Measurements of the viscosity of 40% glycerol aqueous 5 solution at 23.5°C

The glycerol aqueous solution (2 liters, 40% (w/w)) was prepared by weighing the 6 required amount of 99.5% glycerol (Shanghai Aladdin Biochemical Technology Co., 7 LTD) and diluting it with distilled water. To measure the viscosity, this solution was 8 firstly sampled and poured into a calibrated 0.4 mm capillary viscometer (with an 9 instrument constant  $c = 0.005392 \text{ mm}^2/\text{s}^2$ ), which can measure liquid sample viscosity 10 by measuring fall time driven by gravity in capillary. Then the solution was sent to the 11 phantom for viscosity measurement by the MRI and PBV estimation. At the end, it was 12 submerged in a water bath (23.5°C). 13

14 When the solution viscosity measured by capillary viscometer, the density  $\rho$  was also 15 measured by using a glass densitometer (Weier Instrument & Meter Factory) to be 1120 16 kg·m<sup>-3</sup>.

The fall time t was recorded by a stopwatch as 575.2 s, 587.1 s, and 578.1 s in three
independent capillary viscometer tests, respectively. The corresponding viscosities
were 3.470 mPa·s, 3.541 mPa·s, and 3.498 mPa·s (calculated using the Formula S1).
The average value of these viscosities is 3.503 mPa·s.

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$$\mu (Pa \cdot s) = \frac{c (mm^2 \cdot s^{-2}) \cdot t (s) \cdot \rho (kg \cdot m^{-3})}{10^6 (mm^2 \cdot m^{-2})}$$
(S1)

### 22 Section 2: Supplementary Figures & Tables

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Group	Rat No.	Day 0	Day 10	Day 22	Day 31	Day 41
	1	225.68	265.96	558.39	878.76	553.90
	2	630.13	614.36	1031.24	759.13	409.44
HLP	3	263.88	564.42	468.41	835.62	785.03
	4	213.16	455.64	889.23	563.17	730.54
	5	520.11	171.17	828.42	847.99	555.83
	6	762.14	1005.11	604.13	727.69	590.46
	7	193.34	673.94	670.13	655.01	915.87
NC	8	349.18	670.88	929.61	642.29	300.90
	9	622.37	305.80	728.56	583.48	739.41
	10	534.59	482.74	841.93	578.94	851.47

\*The maximum value of Reynold's number among common, internal and external carotid artery in each rat is shown in this table. If the Reynold's number Re < 2000, the blood flow is laminar under the corresponding conditions.

	Parameter name	Cohen's d (t-test)/ Effect size f (rmANOVA)	Minimum sample size per group (alpha=0.05, 80% power)	Actual sample size per group
	LDL-C (rat)	3.85	3	5
	LDL-C (rabbit)	7.01	3	3
	TAWSS <sub>min</sub> (Day 31)	2.77	3	5
	OSI <sub>max</sub> (Day 42)	1.90	5	5
	Low WSS level (W4)	3.43	3	5
	LCCA diameter (W7)	1.56	7	7
	Hematocrit (rat)	2.04	4	5
	Diet (rat, rmANOVA)	0.70	3	5
	Diet×PBV (rat, rmANOVA)	0.93	3	5
	Diet (rabbit, rmANOVA)	0.37	5	5
	Diet×PBV (rabbit, rmANOVA)	0.48	4	5

 Table S2 Sample size for each parameter (post-hoc)
 Image: size for each parameter (post-hoc)

\*LDL-C: low-density lipoprotein cholesterol, TAWSS: time-averaged wall shear
 stress, OSI: Oscillatory shear index, WSS: wall shear stress, LCCA: left common
 carotid artery, PBV: personalized blood viscosity.





Fig.S1 Blood lipid profiles in rats (Day 42, n=5) and rabbits (Week 4, n=3). (a) HLPrat group showed elevated TC and LDL-C compared to NC-rat group. (b) HLP-rabbit
group exhibited combined hyperlipidemia. \*: p<0.05, \*\*: p<0.01 (unpaired t-test).</li>



- 41 Fig.S2 Oscillatory Shear Index (OSI) maps in rats. HLP2, HLP4, and HLP5 columns
- 42 are three series of OSI maps from the HLP-rat group, while the maps in NC7 column
- 43 are from NC-rat group. The HLP5\_emp column shows OSI results using the empirical
- 44 power-law blood viscosity model. Arrows indicate regions of high OSI (>0.3).



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Fig.S3 Temporal evolution of time-average wall shear stress (TAWSS) and oscillatory 46 shear index (OSI) in rats (a) Minimum TAWSS values in HLP-rat and NC-rat groups 47 with PBV correction. (b) Maximum OSI values. Data presented as mean  $\pm$  SD. \*: 48 p<0.05, \*\*: p<0.01 (n=5, unpaired t-test). 49



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51 Fig.S4 TAWSS differences between HLP-rat and NC-rat groups on Day 31. PBVcorrected TAWSS values (0.62  $\pm$  0.13 Pa vs. 1.00  $\pm$  0.15 Pa, p<0.05), empirical 52 model TAWSS values (0.79  $\pm$  0.15 Pa vs. 0.83  $\pm$  0.16 Pa, p=0.77). (n=5, unpaired 53 54 t-test).



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Fig. S5 Low WSS area quantification in rabbits. (a) PBV-corrected results showing 56 significant differences between HLP-rabbit and NC-rabbit groups at Week 4 and 57 Week 7. (b) Empirical model results showed no significance. \*: p<0.05 (n=5; 58 unpaired t-test).



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Fig.S6 H&E-stained sections of HLP rabbit (A) and healthy rabbit (B) at week 4
 (scale bar: 200 μm). The magnified image of the smooth muscle proliferation lesion

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Fig.S7 H&E-stained sections of HLP rabbit (A) and healthy rabbit (B) at week 7
(scale bar: 200 μm). The magnified image of the Atherosclerosis lesion is shown in
Figure C (scale bar: 50 μm).



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69 Fig.S8 The typical TOF-MRA image of hyperlipidemia rabbit and healthy rabbit at

- 70 Week 4 and Week 7. The signal of the left external carotid artery highlighted by
- 71 yellow arrows disappeared over time due to the change of vascular geometry.



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Fig.S9 The rabbits' left common carotid arteries diameters at week 0, 4 and 7 in the
 TOF-MRI image. \*\*: p=0.01 (n=7, unpaired t-test).



Fig.S10 Hematocrit results of SD rats (on Day 42) and NZW rabbits (at Week 7) with
 high-fat diet and conventional diet. \*: p<0.05 (n=5, unpaired t-test).</li>