## **Supplementary Materials**



Figure S1. Expression detection of the target receptors on the inflammatory cells stimulated with TNF- $\alpha$ . Expression of VCAM-1, ICAM-1 and P-selectin on bEnd.3 cells which were stimulated by TNF- $\alpha$  with different concentration and incubating time were analyzed by flow cytometry. \*P < 0.05.



Figure S2. In vitro dynamic adhesion of MBs. Representative micrographs after incubating bEnd.3 cells with targeted MBs for 4 min with a flow velocity at 4 dynes/cm<sup>2</sup> shear stress (bar = 50  $\mu$ m).



Figure S3. In vivo results of UMI performed with  $MB_{VIS}$  and control MBs in ascending aorta. Mice from four different groups were detected by UMI at 6-, 10- or 14-week feeding time point, respectively. A good performance of  $MB_{VIS}$  could be discovered. Molecular imaging effect of targeted MBs had a significant enhancement than that of  $MB_{IgG}$  (\*: P<0.05 ), and  $MB_{VIS}$  had better imaging effect than that of the single- and dual-targeted MBs in the most time points and animal groups (\*: P<0.05). The signal intensity of all targeted MBs showed an increasing trend along with feeding time.

Commlo	Number-weighted diameter (µm)			
Sample	Mean ±SD	Median $\pm$ SD	Mode ±SD	
MB <sub>IgG</sub>	$2.34 \pm 0.50$	$1.78 \pm 0.28$	$1.75 \pm 0.06$	
$\mathrm{MB}_{\mathrm{V}}$	$2.05 \pm 0.29$	$1.59 \pm 0.21$	$1.61 \pm 0.22$	
$MB_{I}$	$2.01 \pm 0.17$	$1.62 \pm 0.03$	$1.75 \pm 0.06$	
MBs	$2.07 \pm 0.34$	$1.60 \pm 0.09$	$1.71 \pm 0.06$	
$MB_{VI}$	$2.19 \pm 0.56$	$1.71 \pm 0.06$	$1.75 \pm 0.30$	
$MB_{VS}$	$2.13 \pm 0.17$	$1.63 \pm 0.18$	$1.57 \pm 0.13$	
MB <sub>IS</sub>	$2.33 \pm 0.52$	$1.7\ \pm 0.06$	$1.81 \pm 0.25$	
MB <sub>VIS</sub>	$2.40 \pm 0.45$	$1.63 \pm 0.18$	$1.69 \pm 0.18$	

Table 1. Particle size of the targeted MBs and control isotype MBs.

Table 2. The ligand amount conjugated onto the microbubble surface.

	VCAM-1 antibody	ICAM-1 antibody	Sialyl Lewis-X polymer
	( $10^{5/}$ microbubble )	( $10^{5/}$ microbubble )	( 10 <sup>5</sup> / microbubble )
$MB_V$	12.9±0.87		
$MB_{I}$		13.68±0.68	
MBs			2.25±0.19
$MB_{VI}$	5.71±0.21	6.98±0.36	
$MB_{VS}$	4.71±0.18		$0.91 \pm 0.04$
MB <sub>IS</sub>		6.31±0.73	1.29±0.40
MB <sub>VIS</sub>	4.26±0.27	5.51±0.29	$0.93 \pm 0.04$

Note: "--" means that the ligand amount was not detected.

ApoE <sup>-/-</sup> HD mice		HDLC(mmol/L)	LDLC(mmol/L)	TG(mmol/L)	T-CHO(mmol/L)
Control	6 weeks	$1.4 \pm 0.55$	$1.15\ \pm 0.26$	$1.36 \pm 0.44$	12.52 ±2.95
Ator	10 weeks	2.13 ±1.13	$2.88 \pm 1.38$	$2.67 \pm 1.72$	22.21 ±3.31 <sup>#</sup>
	14 weeks	$2.45 \pm 0.61^{\#}$	$2.62 \pm 0.57*$	4.27 ±0.48*#	15.31 ±3.95 <sup>#</sup>
Placebo	10 weeks	2.13 ±1.13	$2.88 \pm 1.38$	$2.67 \pm 1.72$	22.21 ±3.31
	14 weeks	6.34 ±0.97*	5.71 ±1.92*	6.61 ±1.43*	25.53 ±5.15*

Table S3. Serum lipid profile of A-HD mice under atorvastatin or placebo treatment.

Note: HDLC is for high density lipoprotein cholesterol, LDLC for low density lipoprotein cholesterol, TG for triglyceride, T-CHO for total cholesterol. They were measured by enzymatic assays. \*: P<0.05, compare with the serum lipid value of six-week-feeding A-HD mice. #: P<0.05, compare with the serum lipid value of placebo treated groups.