Supplementary information for

Local Production of Activated Factor X in Atherosclerotic

Plaque Induced Vascular Smooth Muscle Cell Senescence

Fumihiro Sanada¹, Jun Muratsu^{1, 2}, Rei Otsu¹, Hideo Shimizu¹, Nobutaka Koibuchi³,

Kazutaka Uchida⁴, Yoshiaki Taniyama^{1, 2}, Shinichi Yoshimura⁴, Hiromi Rakugi², Ryuichi

Morishita¹

¹Department of Clinical Gene Therapy, ²Department of Geriatric and General Medicine,

Osaka University Graduate School of Medicine, Suita, Osaka 565-0871, Japan

³Departments of Pharmacology and Molecular Therapeutics, Kumamoto University

Graduate School of Medical Sciences, Kumamoto, Japan

⁴Department of Neurosurgery, Hyogo College of Medicine, Hyogo, Japan

Running title

FXa induces smooth muscle cell senescence.

Address correspondence to:

Yoshiaki Taniyama, MD, PhD, Associate Professor

Department of Clinical Gene Therapy, Osaka University Graduate School of Medicine.

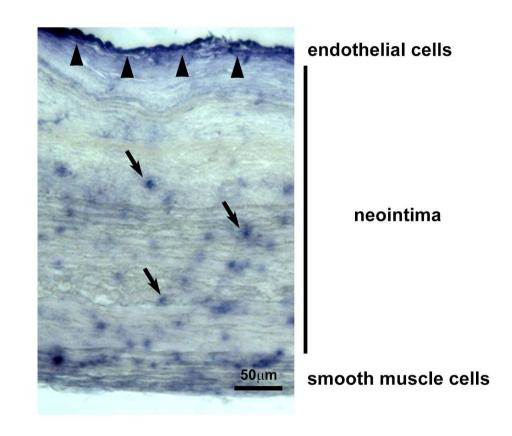
2-2 Yamada-oka, Suita, Osaka, 565-0871

Tel.: +81-6-6210-8351, FAX: +81-6-6210-8359

E-mail address; taniyama@cgt.med.osaka-u.ac.jp

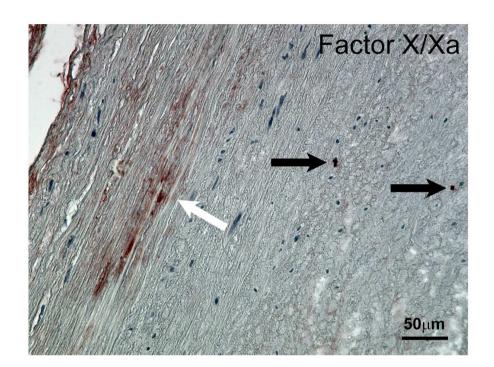
Ryuichi Morishita, MD, PhD, Professor

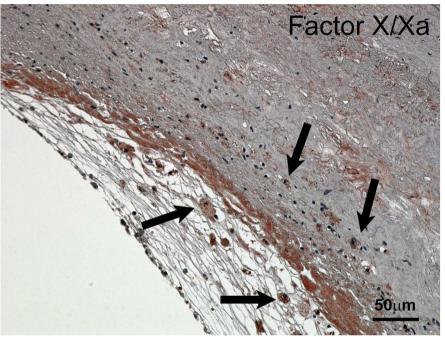
E-mail: morishit@cgt.med.osaka-u.



Supplementary figure 1

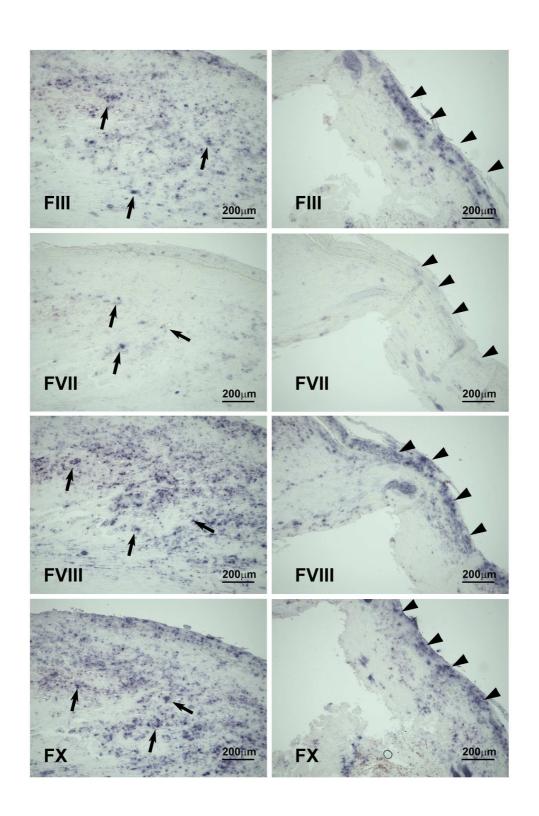
Factor X expression in human atherosclerotic plaques. Strong FX mRNA signal was detected in VSMCs, endothelial cells (arrow head) and inflammatory cells (black arrow).



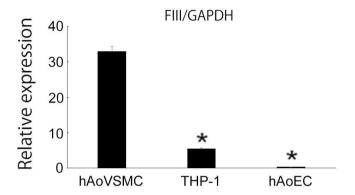


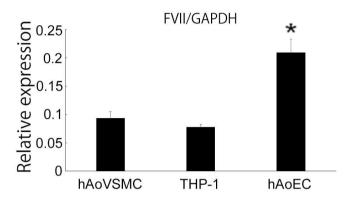
Supplementary figure 2

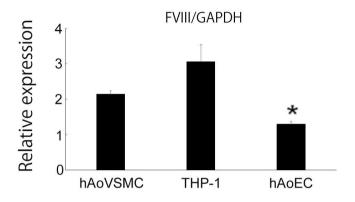
Factor X/Xa expression in human atherosclerotic plaques. Strong FX/FXa signals were detected in VSMCs (white arrow) and inflammatory cells (black arrow).

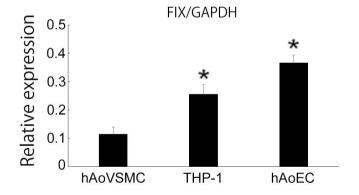


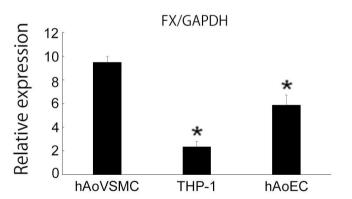
Supplementary figure 3
Factor III, VII, VIII, and X mRNA expression in human atherosclerotic plaques. Strong FIII, FVIII, and FX mRNA signals was detected in inflammatory cells (arrow head) and VSMCs (black arrow). FVII signal was relatively weak, but detected in VSMCs and inflammatory cells..



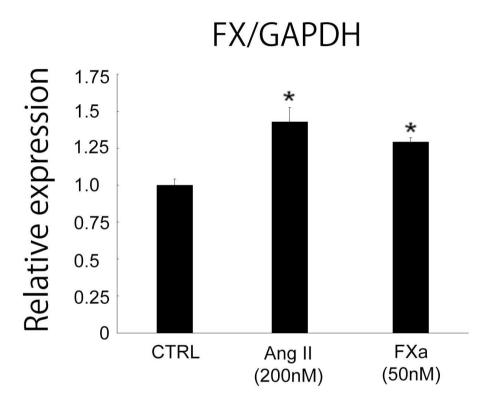








Supplementary figure 4 FIII, FVII, FVIII, FIX, and FX mRNA expression in hAoVSMC, THP-1, and hAoEC. *P<0.05 vs. hAoVSMC, n=5.



Supplementary figure 5 FX mRNA expression in human aortic VSMCs in vitro. Ang II = angiotensin II. *P<0.05 vs. control, n=6.