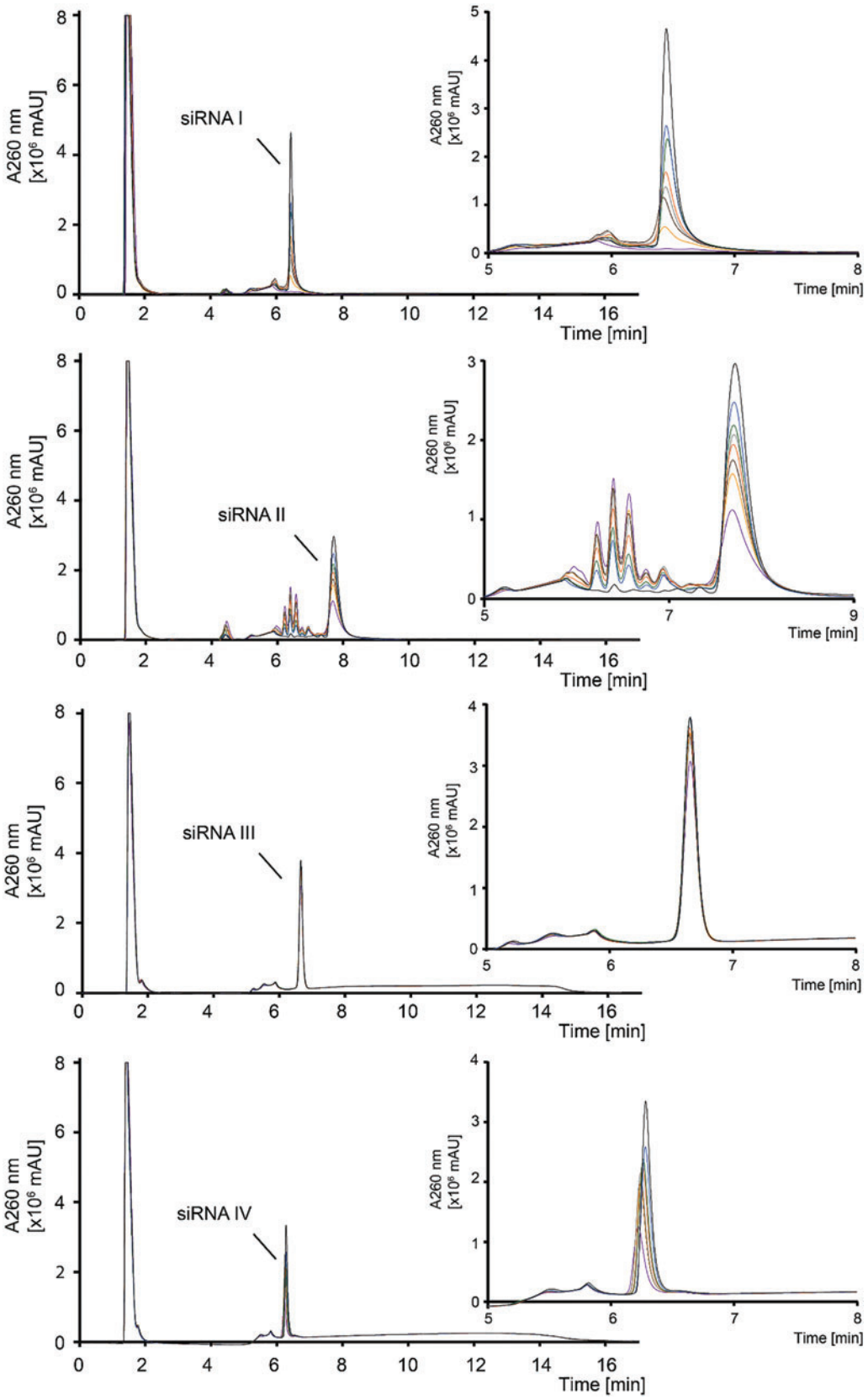
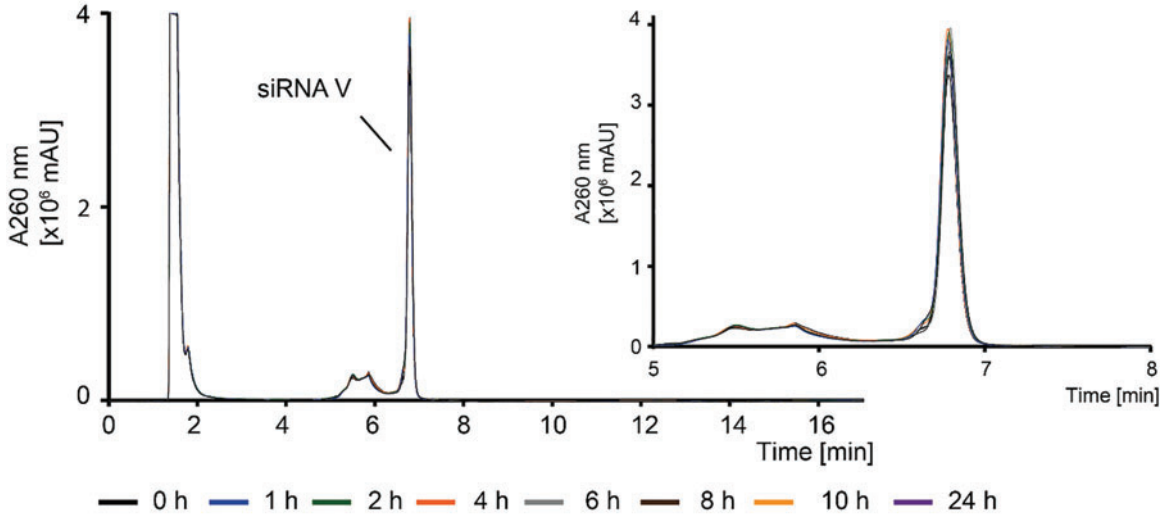


SUPPLEMENTARY FIG. S3. Representative anion-exchange HPLC chromatograms from serum stability assays. siRNAs were incubated in 50% mouse serum for the indicated time points. Samples were analyzed on a Hitachi VWR LaChrom Elite HPLC fitted with a DNA Pac PA200 (4×250 mm) anion exchange column and a DNA Pac PA200 (4×50 mm) guard column at 30°C. UV absorption was monitored at 260 nm. The gradient was 100% A for 2 min, followed by 54% eluent B within 5 min, increase to 100% B within 2 min, hold 100% B for 1 min, switch to 100% A within 2 min, and hold 100% A for 5 min. Eluent A was a 1:1 mixture (v:v) of buffer A and ACN. Buffer A was an aqueous solution of 1 mM EDTA and 25 mM Tris HCl (pH = 8.5). Eluent B was a 1:1 mixture (v:v) of buffer B and ACN. Buffer B was an aqueous solution of 1 mM EDTA, 25 mM Tris HCl, and 1.6 M $NaClO_4$ (pH = 8.5). HPLC, high performance liquid chromatography; EDTA, ethylenediaminetetraaceticacid; ACN, acetonitrile.

(*continued*)



SUPPLEMENTARY FIG. S3. (*Continued*).