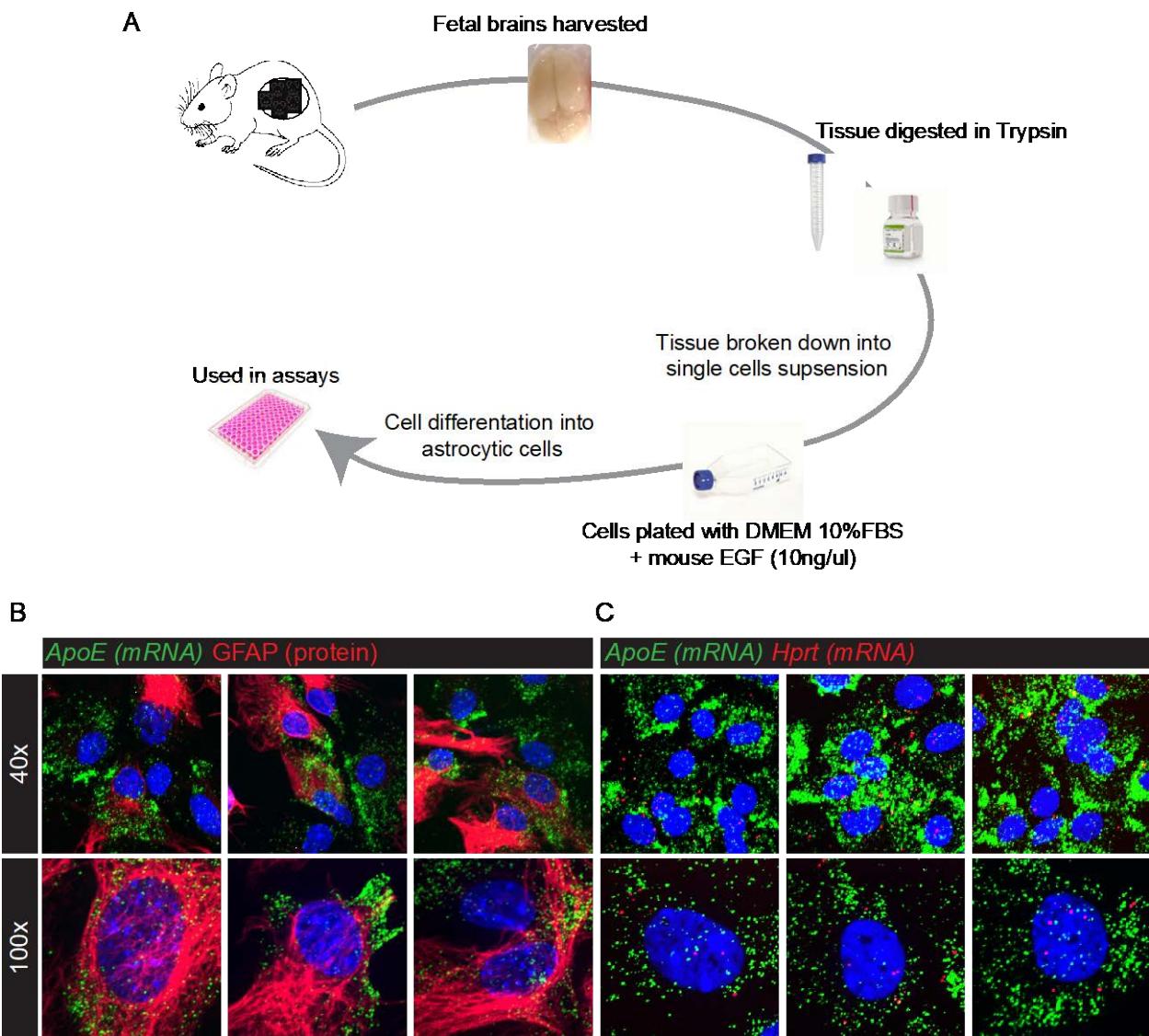


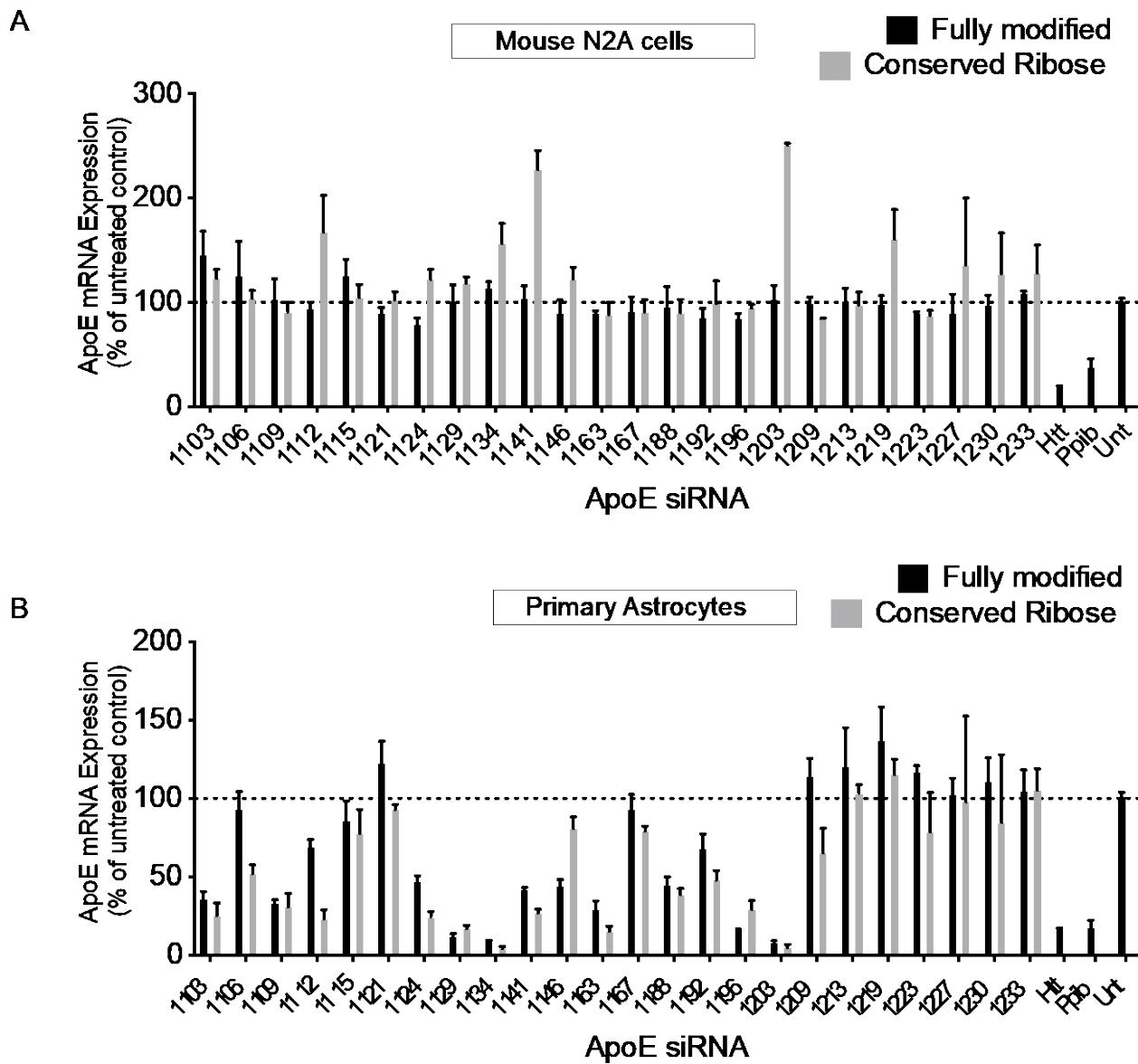
Supplemental Information

**Cell Type Impacts Accessibility of mRNA
to Silencing by RNA Interference**

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Supplementary Figure 1: Preparation of primary mouse astrocytes. (A) Workflow for harvest and preparation. (B) Primary mouse astrocytes co-stained for *ApoE* mRNA (green) (RNAscope) and GFAP protein (red) (IF) at 40x and 100x magnification confirming astrocyte lineage. (C) Primary mouse astrocytes co-stained for *ApoE* (green) and *HPRT* (red) mRNA (RNAscope) at 40x and 100x magnification.



Supplementary Figure 2: Conservation of 2'OH (Ribose) does not impact RNAi efficacy.

(A). Screen using fully modified (black) and partially modified (grey) siRNAs targeting *ApoE* in N2A cells. (B) Screen using fully modified (black) and partially modified (grey) siRNAs targeting *ApoE* in mouse primary astrocytes. Dose: 1.5 μ M; 72 -hour timepoint; mRNA quantified using QuantiGene.

Table 1: siRNA sequences and chemical modifications

Name	Gene	Accession Number	Anti-sense strand	Sense strand
Apoe_156	Mouse ApoE	NM_009696	P(mU)##(fC)##(mA)(fA)(mU)(fU)(mG)(fU)(mG)(fA)(mU)(fU)(mG)##(fG)##(mC)##(fC)##(mA)##(fG)##(mU)##(fC)	(fG)##(mC)##(fC)(mA)(fA)(mU)(fC)(mA)(fC)(mA)(fA)(mU)(fU)##(mG)##(fA)-TegChol
Apoe_416	Mouse ApoE	NM_009696	P(mU)##(fU)##(mU)(fU)(mG)(fU)(mA)(fA)(mG)(fC)(mC)(fU)(mU)##(fU)##(mA)##(fC)##(mU)##(fU)##(mC)##(fC)	(fU)##(mA)##(fA)(mA)(fG)(mG)(fC)(mU)(fU)(mA)(fC)(mA)(fA)##(mA)##(fA)-TegChol
Apoe_636	Mouse ApoE	NM_009696	P(mU)##(fU)##(mC)(fA)(mA)(fG)(mC)(fG)(mC)(fU)(mU)(fG)(mC)##(fG)##(mC)##(fA)##(mU)##(fC)##(mU)##(fU)	(fG)##(mC)##(fG)(mC)(fA)(mA)(fG)(mC)(fC)(mC)(fU)(mU)(fG)##(mA)##(fA)-TegChol
Apoe_1004	Mouse ApoE	NM_009696	P(mU)##(fG)##(mU)(fC)(mU)(fU)(mC)(fC)(mA)(fC)(mU)(fA)(mU)##(fU)##(mG)##(fG)##(mC)##(fU)##(mC)##(fG)	(fC)##(mA)##(fA)(mU)(fA)(mG)(fU)(mG)(fG)(mA)(fA)(mG)(fA)##(mC)##(fA)-TegChol
Apoe_1032	Mouse ApoE	NM_009696	P(mU)##(fC)##(mC)(fA)(mU)(fC)(mA)(fG)(mG)(fU)(mU)(fU)(mG)##(fC)##(mC)##(fC)##(mA)##(fC)##(mU)##(fG)	(fG)##(mG)##(fC)(mA)(fA)(mA)(fC)(mA)(fC)(fU)(mG)(fA)(mU)(fG)##(mG)##(fA)-TegChol
Apoe_1103	Mouse ApoE	NM_009696	P(mU)##(fG)##(mG)(fA)(mU)(fA)(mC)(fU)(mC)(fA)(mU)(fU)(mG)##(fA)##(mU)##(fU)##(mC)##(fU)##(mC)##(fC)	(fA)##(mU)##(fC)(mA)(fA)(mU)(fG)(mA)(fG)(mU)(fA)(mU)(fC)##(mC)##(fA)-TegChol
Apoe_1106	Mouse ApoE	NM_009696	P(mU)##(fG)##(mA)(fA)(mG)(fG)(mA)(fU)(mA)(fC)(mU)(fC)(mA)##(fU)##(mU)##(fG)##(mA)##(fU)##(mU)##(fC)	(fA)##(mA)##(fU)(mG)(fA)(mG)(fU)(mA)(fU)(mC)(fC)(mU)(fU)##(mC)##(fA)-TegChol
Apoe_1109	Mouse ApoE	NM_009696	P(mU)##(fG)##(mG)(fA)(mG)(fA)(mA)(fG)(mG)(fA)(mU)(fA)(mC)##(fU)##(mC)##(fA)##(mU)##(fU)##(mG)##(fA)	(fG)##(mA)##(fG)(mU)(fA)(mU)(fC)(mC)(fU)(mU)(fC)##(mC)##(fA)-TegChol
Apoe_1112	Mouse ApoE	NM_009696	P(mU)##(fA)##(mC)(fA)(mG)(fG)(mA)(fG)(mA)(fA)(mG)(fG)(mA)##(fU)##(mA)##(fC)##(mU)##(fC)##(mA)##(fU)	(fU)##(mA)##(fU)(mC)(fC)(mU)(fU)(mC)(fC)(mU)(fC)##(mU)##(fA)-TegChol
Apoe_1115	Mouse ApoE	NM_009696	P(mU)##(fA)##(mG)(fG)(mA)(fC)(mA)(fG)(mG)(fA)(mG)(fA)(mA)##(fG)##(mG)##(fA)##(fA)##(mU)##(fC)##(mA)##(fU)	(fC)##(mC)##(fU)(mU)(fC)(mU)(fC)(mC)(fU)(mG)(fU)(mC)(fC)##(mU)##(fA)-TegChol
Apoe_1119	Mouse ApoE	NM_009696	P(mU)##(fU)##(mU)(fG)(mC)(fA)(mG)(fG)(mA)(fC)(mA)(fG)(mG)##(fA)##(mG)##(fA)##(fG)##(mG)##(fA)	(fC)##(mU)##(fC)(mC)(fU)(mG)(fU)(mC)(fC)(mU)(fG)(mC)(fA)##(mA)##(fA)-TegChol
Apoe_1121	Mouse ApoE	NM_009696	P(mU)##(fU)##(mG)(fU)(mU)(fG)(mC)(fA)(mG)(fG)(mA)##(fC)##(mG)##(fA)##(fG)##(fA)##(mA)##(fG)	(fC)##(mC)##(fU)(mG)(fU)(mC)(fC)(mU)(fG)(mC)(fA)##(mA)##(fA)-TegChol
Apoe_1123	Mouse ApoE	NM_009696	P(mU)##(fG)##(mU)(fU)(mG)(fU)(mU)(fG)(mC)(fA)(mG)(fG)(mA)##(fC)##(mA)##(fG)##(fA)##(mG)##(fA)##(fG)	(fU)##(mG)##(fU)(mC)(fC)(mU)(fG)(mC)(fA)(mA)(fC)(mA)##(mC)##(fA)-TegChol
Apoe_1124	Mouse ApoE	NM_009696	P(mU)##(fU)##(mG)(fU)(mU)(fG)(mU)(fU)(mG)(fC)(mA)(fG)(mG)##(fA)##(mC)##(fA)##(fG)##(fG)##(mA)##(fG)	(fG)##(mU)##(fC)(mC)(fU)(mG)(fC)(mA)(fA)(mA)(fC)##(mA)##(fA)-TegChol
Apoe_1126	Mouse ApoE	NM_009696	P(mU)##(fG)##(mA)(fU)(mG)(fU)(mU)(fG)(mU)(fU)(mG)(fC)(mA)##(fG)##(mG)##(fA)##(fC)##(fC)##(mA)##(fG)##(fG)	(fC)##(mC)##(fU)(mG)(fC)(mA)(fA)(mA)(fC)(mA)(fA)(fC)##(mA)##(fC)-TegChol
Apoe_1129	Mouse ApoE	NM_009696	P(mU)##(fA)##(mU)(fG)(mG)(fA)(mU)(fG)(mU)(fU)(mG)(fU)(mU)##(fG)##(mC)##(fA)##(fG)##(fG)##(mA)##(fC)	(fG)##(mC)##(fA)(mA)(fC)(mA)(fA)(mA)(fC)(mA)(fC)(fA)##(mA)##(fA)-TegChol
Apoe_1134	Mouse ApoE	NM_009696	P(mU)##(fU)##(mG)(fG)(mA)(fU)(mA)(fU)(mG)(fG)(mA)(fU)(mG)##(fU)##(mA)##(fG)##(fU)##(mU)##(fG)##(fC)##(fC)	(fA)##(mA)##(fC)(mA)(fU)(mC)(fC)(mA)(fU)(mA)(fC)(mA)(fC)##(mA)##(fA)-TegChol
Apoe_1141	Mouse ApoE	NM_009696	P(mU)##(fC)##(mC)(fC)(mU)(fG)(mG)(fC)(mU)(fG)(mG)(fA)(mU)##(fA)##(mU)##(fG)##(fA)##(fG)##(fA)##(fG)	(fA)##(mU)##(fA)(mA)(fC)(mC)(fA)(mA)(fC)(mA)(fG)##(mU)##(fA)-TegChol
Apoe_1146	Mouse ApoE	NM_009696	P(mU)##(fG)##(mG)(fG)(mC)(fC)(mA)(fC)(mC)(fU)(mG)(fG)(mC)##(fU)##(fG)##(fG)##(mA)##(fU)##(fA)##(fG)	(fC)##(mA)##(fG)(mC)(fC)(mA)(fG)(mG)(fU)(mG)(fG)(mC)(fC)##(mA)##(fA)-TegChol
Apoe_1163	Mouse ApoE	NM_009696	P(mU)##(fG)##(mA)(fG)(mG)(fG)(mU)(fG)(mG)(fC)(mU)(fU)(mG)##(fA)##(mG)##(fA)##(fC)##(fC)##(mA)##(fG)##(fG)	(fC)##(mU)##(fC)(mA)(fA)(mA)(fG)(fC)(mA)(fC)(mA)(fC)##(mA)##(fA)-TegChol
Apoe_1167	Mouse ApoE	NM_009696	P(mU)##(fG)##(mC)(fC)(mA)(fG)(mA)(fG)(mA)(fG)(mG)(fG)(mU)(fG)##(fC)##(mU)##(fU)##(mG)##(fA)##(fG)##(fA)	(fA)##(mG)##(fC)(mA)(fC)(mC)(fU)(mC)(fC)(mA)(fG)##(mC)##(fA)-TegChol

Apoe_1188	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mU)(fA)(mA)(fG)(mC)(fA)(mA)(fG)(mG)(fG)(mC)#{(fC)#{(mA)#{(fC)#{(mC)#{(fA)#{(mG)#{(fA)}}	(fU)#{(mG)#{(fG)(mC)(fC)(mC)(fU)(mU)(fG)(mC)(fU)(mU)(fA)#{(mA)#{(fA)-TegChol
Apoe_1191	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mU)(fA)(mU)(fU)(mA)(fA)(mG)(fC)(mA)(fA)(mG)#{(fG)#{(mG)#{(fC)#{(mC)#{(fA)#{(mC)#{(fC)}}	(fC)#{(mC)#{(fC)(mU)(fU)(mG)(fC)(mU)(fU)(mA)(fA)(mU)(fA)#{(mA)#{(fA)-TegChol
Apoe_1192	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mU)(fU)(mA)(fU)(mU)(fA)(mA)(fG)(mC)(fA)(mA)#{(fG)#{(mG)#{(fG)#{(mC)#{(fC)#{(mA)#{(fC)}}	(fC)#{(mC)#{(fU)(mU)(fG)(mC)(fU)(mU)(fA)(mA)(fA)#{(mA)#{(fA)-TegChol
Apoe_1196	Mouse ApoE	NM_009696	P(mU)#{(fA)#{(mA)(fU)(mC)(fU)(mU)(fU)(mA)(fU)(mU)(fA)(mA)#{(fG)#{(mC)#{(fA)#{(mA)#{(fG)#{(mG)#{(fG)}}	(fG)#{(mC)#{(fU)(mU)(fA)(mA)(fU)(mA)(fA)(mA)(fG)(mA)(fA)(fU)#{(mU)#{(fA)-TegChol
Apoe_1203	Mouse ApoE	NM_009696	P(mU)#{(fC)#{(mU)(fC)(mG)(fG)(mA)(fG)(mA)(fA)(mU)(fC)(mU)#{(fU)#{(mA)#{(fA)#{(mU)#{(fU)#{(mA)#{(fA)}}	(fA)#{(mA)#{(fA)(mG)(fA)(mU)(fU)(mC)(fU)(mC)(fA)#{(mG)#{(fA)-TegChol
Apoe_1209	Mouse ApoE	NM_009696	P(mU)#{(fA)#{(mA)(fU)(mG)(fU)(mG)(fC)(mU)(fC)(mG)(fG)(mA)#{(fG)#{(mA)#{(fA)#{(mU)#{(fC)#{(mU)#{(fU)}}	(fU)#{(mC)#{(fU)(mC)(fC)(mG)(fA)(mG)(fC)(mA)(fC)(mA)(fU)#{(mU)#{(fA)-TegChol
Apoe_1213	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mC)(fA)(mG)(fA)(mA)(fU)(mG)(fU)(mG)(fC)(mU)#{(fC)#{(mG)#{(fG)#{(mA)#{(fG)#{(mA)#{(fA)}}	(fC)#{(mG)#{(fA)(mG)(fC)(mA)(fC)(mA)(fU)(mU)(fC)(mU)(fG)#{(mA)#{(fA)-TegChol
Apoe_1213	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mC)(fA)(mG)(fA)(mA)(fU)(mG)(fU)(mG)(fC)(mU)#{(fC)#{(mG)#{(fG)#{(mA)#{(fG)#{(mA)#{(fA)}}	(fC)#{(mG)#{(fA)(mG)(fC)(mA)(fC)(mA)(fU)(mU)(fC)(mU)(fG)#{(mA)#{(fA)-TegChol
Apoe_1217	Mouse ApoE	NM_009696	P(mU)#{(fA)#{(mG)(fA)(mC)(fU)(mC)(fA)(mG)(fA)(mA)(fU)(mG)#{(fU)#{(mG)#{(fC)#{(mU)#{(fC)#{(mG)#{(fG)}}	(fC)#{(mA)#{(fC)(mA)(fU)(mU)(fC)(mU)(fG)(mA)(fG)(mU)(fC)#{(mU)#{(fA)-TegChol
Apoe_1219	Mouse ApoE	NM_009696	P(mU)#{(fA)#{(mG)(fA)(mG)(fA)(mC)(fU)(mC)(fA)(mG)(fA)(mA)#{(fU)#{(mG)#{(fU)#{(mG)#{(fC)#{(mU)#{(fC)}}	(fC)#{(mA)#{(fU)(mU)(fC)(mU)(fG)(mA)(fG)(mU)(fC)#{(mU)#{(fA)-TegChol
Apoe_1223	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mC)(fA)(mC)(fA)(mG)(fA)(mA)(fG)(fA)(mC)(fU)#{(fA)#{(mG)#{(fA)#{(mA)#{(fU)#{(mG)#{(fU)}}	(fC)#{(mU)#{(fG)(mA)(fG)(mU)(fC)(mU)(fC)(mU)(fG)#{(mA)#{(fA)-TegChol
Apoe_1227	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mC)(fA)(mC)(fU)(mC)(fA)(mC)(fA)(mG)(fA)(mA)#{(fA)#{(mC)#{(fU)#{(mC)#{(fA)#{(fA)#{(mG)#{(fA)}}	(fG)#{(mU)#{(fC)(mU)(fC)(mU)(fG)(mA)(fG)(mU)(fG)(mA)(fG)(mU)(fG)#{(mA)#{(fA)-TegChol
Apoe_1230	Mouse ApoE	NM_009696	P(mU)#{(fG)#{(mA)(fA)(mU)(fC)(mA)(fC)(mU)(fC)(mA)(fC)(mA)#{(fG)#{(mA)#{(fG)#{(mA)#{(fC)#{(mU)#{(fC)}}	(fU)#{(mC)#{(fU)(mG)(fU)(mG)(fA)(mG)(fU)(mG)(fA)(mU)(fU)#{(mC)#{(fA)-TegChol
Apoe_1233	Mouse ApoE	NM_009696	P(mU)#{(fU)#{(mU)(fG)(mG)(fA)(mA)(fU)(mC)(fA)(mC)(fU)(mC)#{(fA)#{(mC)#{(fA)#{(mG)#{(fA)#{(fA)#{(mG)#{(fA)}}	(fG)#{(mU)#{(fG)(mA)(fG)(mU)(fG)(mA)(fU)(mU)(fC)(mC)(fA)#{(mA)#{(fA)-TegChol

Detailed sequence and chemical modification patterns, of siRNAs. Chemical modifications are designated as follows, “#” – phosphorothioate bond, “m” – 2'-O-Methyl, “f” – 2'-Fluoro, “P” – 5' Phosphate.

Table 2: Partially modified sense strand sequences and chemical pattern

Name	Gene	Accession Number	Sense strand
Apoe_156	Mouse ApoE	NM_009696	(fA)#{(mU)#{(fC)(rA)(rA)(rU)(fG)(mA)(fG)(mU)(fA)(mU)(fC)#{(mC)#{(fA)-TegChol}}
Apoe_416	Mouse ApoE	NM_009696	(fA)#{(mA)#{(fU)(rG)(rA)(rG)(fU)(mA)(fU)(mC)(fC)(mU)(fU)#{(mC)#{(fA)-TegChol}}
Apoe_636	Mouse ApoE	NM_009696	(fG)#{(mA)#{(fG)(rU)(rA)(rU)(fC)(mC)(fU)(mU)(fC)(mC)#{(mC)#{(fA)-TegChol}}
Apoe_1004	Mouse ApoE	NM_009696	(fU)#{(mA)#{(fU)(rC)(rC)(rU)(fU)(mA)(fC)(fC)(mU)(fG)#{(mU)#{(fA)-TegChol}}
Apoe_1032	Mouse ApoE	NM_009696	(fC)#{(mC)#{(fU)(rU)(rC)(fC)(mC)(fU)(mG)(fU)(mC)(fC)#{(mU)#{(fA)-TegChol}}
Apoe_1103	Mouse ApoE	NM_009696	(fC)#{(mC)#{(fU)(rG)(rU)(rC)(fC)(mU)(fG)(mC)(fA)(mA)#{(mA)#{(fA)-TegChol}}
Apoe_1106	Mouse ApoE	NM_009696	(fG)#{(mU)#{(fC)(rC)(rU)(rG)(fC)(mA)(fA)(mC)(fA)(mA)#{(mA)#{(fA)-TegChol}}
Apoe_1109	Mouse ApoE	NM_009696	(fG)#{(mC)#{(fA)(rA)(rC)(rA)(fA)(mA)(fA)(mU)(fC)(mC)(fA)#{(mU)#{(fA)-TegChol}}
Apoe_1112	Mouse ApoE	NM_009696	(fA)#{(mA)#{(fC)(rA)(rU)(rC)(fC)(mA)(fU)(mA)(fC)(mC)#{(mA)#{(fA)-TegChol}}
Apoe_1115	Mouse ApoE	NM_009696	(fA)#{(mU)#{(fA)(rU)(rC)(rC)(fA)(mG)(fC)(mC)(fA)(mG)(fG)#{(mU)#{(fA)-TegChol}}
Apoe_1119	Mouse ApoE	NM_009696	(fC)#{(mA)#{(fG)(rC)(rC)(rA)(fG)(mG)(fU)(mG)(fG)(mC)(fC)#{(mC)#{(fA)-TegChol}}
Apoe_1121	Mouse ApoE	NM_009696	(fC)#{(mU)#{(fC)(rA)(rA)(rG)(fC)(mA)(fC)(mC)(fU)#{(mC)#{(fA)-TegChol}}
Apoe_1123	Mouse ApoE	NM_009696	(fA)#{(mG)#{(fC)(rA)(rC)(rC)(fU)(mC)(fU)(mC)(fU)(mG)(fG)#{(mC)#{(fA)-TegChol}}
Apoe_1124	Mouse ApoE	NM_009696	(fU)#{(mG)#{(fG)(rC)(rC)(fU)(mU)(fG)(mC)(fU)(mU)(fA)#{(mA)#{(fA)-TegChol}}
Apoe_1126	Mouse ApoE	NM_009696	(fC)#{(mC)#{(fU)(rU)(rG)(rC)(fU)(mU)(fA)(mA)(fU)(mA)#{(mA)#{(fA)-TegChol}}
Apoe_1129	Mouse ApoE	NM_009696	(fG)#{(mC)#{(fU)(rU)(rA)(rA)(fU)(mA)(fA)(mA)(fG)(mA)(fU)#{(mU)#{(fA)-TegChol}}
Apoe_1134	Mouse ApoE	NM_009696	(fA)#{(mA)#{(fA)(rG)(rA)(rU)(fU)(mA)(fC)(fG)(mG)#{(mA)#{(fA)-TegChol}}
Apoe_1141	Mouse ApoE	NM_009696	(fU)#{(mC)#{(fU)(rC)(rC)(rG)(fA)(mG)(fC)(mA)(fC)(mA)(fU)#{(mU)#{(fA)-TegChol}}
Apoe_1146	Mouse ApoE	NM_009696	(fC)#{(mG)#{(fA)(rG)(rC)(rA)(fC)(mA)(fU)(mU)(fC)(mU)(fG)#{(mA)#{(fA)-TegChol}}
Apoe_1163	Mouse ApoE	NM_009696	(fC)#{(mA)#{(fU)(rU)(rC)(rU)(fG)(mA)(fG)(mU)(fC)(mU)(fC)#{(mU)#{(fA)-TegChol}}
Apoe_1167	Mouse ApoE	NM_009696	(fC)#{(mU)#{(fG)(rA)(rG)(rU)(fC)(mU)(fC)(mU)(fG)(mU)(fG)#{(mA)#{(fA)-TegChol}}
Apoe_1188	Mouse ApoE	NM_009696	(fG)#{(mU)#{(fC)(rU)(rC)(rU)(fG)(mU)(fG)(mA)(fG)(mU)(fG)#{(mA)#{(fA)-TegChol}}
Apoe_1191	Mouse ApoE	NM_009696	(fU)#{(mC)#{(fU)(rG)(rU)(rG)(fA)(mG)(fU)(mG)(fA)(mU)(fU)#{(mC)#{(fA)-TegChol}}
Apoe_1192	Mouse ApoE	NM_009696	(fG)#{(mU)#{(fG)(rA)(rG)(rU)(fG)(mA)(fU)(mU)(fC)(mC)(fA)#{(mA)#{(fA)-TegChol}}
Apoe_1196	Mouse ApoE	NM_009696	(fA)#{(mU)#{(fC)(rA)(rA)(rU)(fG)(mA)(fG)(mU)(fA)(mU)(fC)#{(mC)#{(fA)-TegChol}}
Apoe_1203	Mouse ApoE	NM_009696	(fA)#{(mA)#{(fU)(rG)(rA)(rG)(fU)(mA)(fU)(mC)(fC)(mU)(fU)#{(mC)#{(fA)-TegChol}}
Apoe_1209	Mouse ApoE	NM_009696	(fG)#{(mA)#{(fG)(rU)(rA)(rU)(fC)(mC)(fU)(mU)(fC)(mU)(fC)#{(mC)#{(fA)-TegChol}}
Apoe_1213	Mouse ApoE	NM_009696	(fU)#{(mA)#{(fU)(rC)(rC)(rU)(fU)(mC)(fU)(mC)(fC)(mU)(fG)#{(mU)#{(fA)-TegChol}}
Apoe_1213	Mouse ApoE	NM_009696	(fC)#{(mC)#{(fU)(rU)(rC)(rU)(fC)(mC)(fU)(mG)(fU)(mC)(fC)#{(mU)#{(fA)-TegChol}}
Apoe_1217	Mouse ApoE	NM_009696	(fC)#{(mC)#{(fU)(rG)(rU)(rC)(fC)(mU)(fG)(mC)(fA)(mA)(fC)#{(mA)#{(fA)-TegChol}}
Apoe_1219	Mouse ApoE	NM_009696	(fG)#{(mU)#{(fC)(rC)(rG)(fC)(mA)(fA)(mC)(fA)(mA)(fC)#{(mA)#{(fA)-TegChol}}
Apoe_1223	Mouse ApoE	NM_009696	(fG)#{(mC)#{(fA)(rA)(rC)(rA)(fA)(mC)(fA)(mU)(fC)(mC)(fA)#{(mU)#{(fA)-TegChol}}
Apoe_1227	Mouse ApoE	NM_009696	(fA)#{(mA)#{(fC)(rU)(rC)(fC)(mA)(fU)(mA)(fC)(mC)(fC)#{(mA)#{(fA)-TegChol}}
Apoe_1230	Mouse ApoE	NM_009696	(fA)#{(mU)#{(fA)(rU)(rC)(rC)(fA)(mG)(fC)(mC)(fA)(mG)(fG)#{(mU)#{(fA)-TegChol}}
Apoe_1233	Mouse ApoE	NM_009696	(fC)#{(mA)#{(fG)(rC)(rA)(fG)(mG)(fU)(mG)(fG)(mC)(fC)#{(mC)#{(fA)-TegChol}}

Detailed sequence and chemical modification patterns, of siRNAs. Chemical modifications are designated as follows, “#” – phosphorothioate bond, “m” – 2'-O-Methyl, “f” – 2'-Fluoro, “P” – 5' Phosphate.

Table 3: Key Resources

Reagent	Source	Identifier
QuantiGene Probeset		
Mouse ApoE	Thermofisher	SB-13611
Mouse PPIB	Thermofisher	SB-10002
Antibodies		
Rabbit polyclonal anti-ApoE	Abcam	183597
Rabbit polyclonal anti-beta-actin	Cell Signaling Technologies	4970
Anti-NeuN	Millipore	MAB37
Anti-GFAP	Abcam	5441
RNAscope Probe sets		
ApoE	ACDBio	313271
HPRT	ACDBio	312951
Commerical Assays		
QuantiGene 2.0	Thermofisher	QS0011
RNAscope Fluorescent Multiplex Kit	ACDBio	320850
Deposited Data		
N2A cells	GSE45119	
Astrocytes	GSE52564	
Primers		
ApoE Forward	GCTCAGACCCTGGAGGCTAA	
ApoE Reverse	CTGTTCCCTCCAGCTCCTTTGTA	