Table S1. Overview of the selected small RNA sequencing kits

Kit	QIAseq miRNA	RealSeq-Biofluids Plasma/Serum miRNA	NEXTFLEX small RNA-seq kit v.3	CleanTag small RNA
Company	Qiagen	RealSeq Biosciences	Perkin Elmer	TriLink Biotechnologies
Basic principle	5’ and 3’ sequential adapter ligation	Single adapter ligation followed by circularization	5’ and 3’ sequential adapter ligation	5’ and 3’ sequential adapter ligation
UMI	+	-	+	-

Figure S1. Read counts distribution in miRXplore and plasma samples

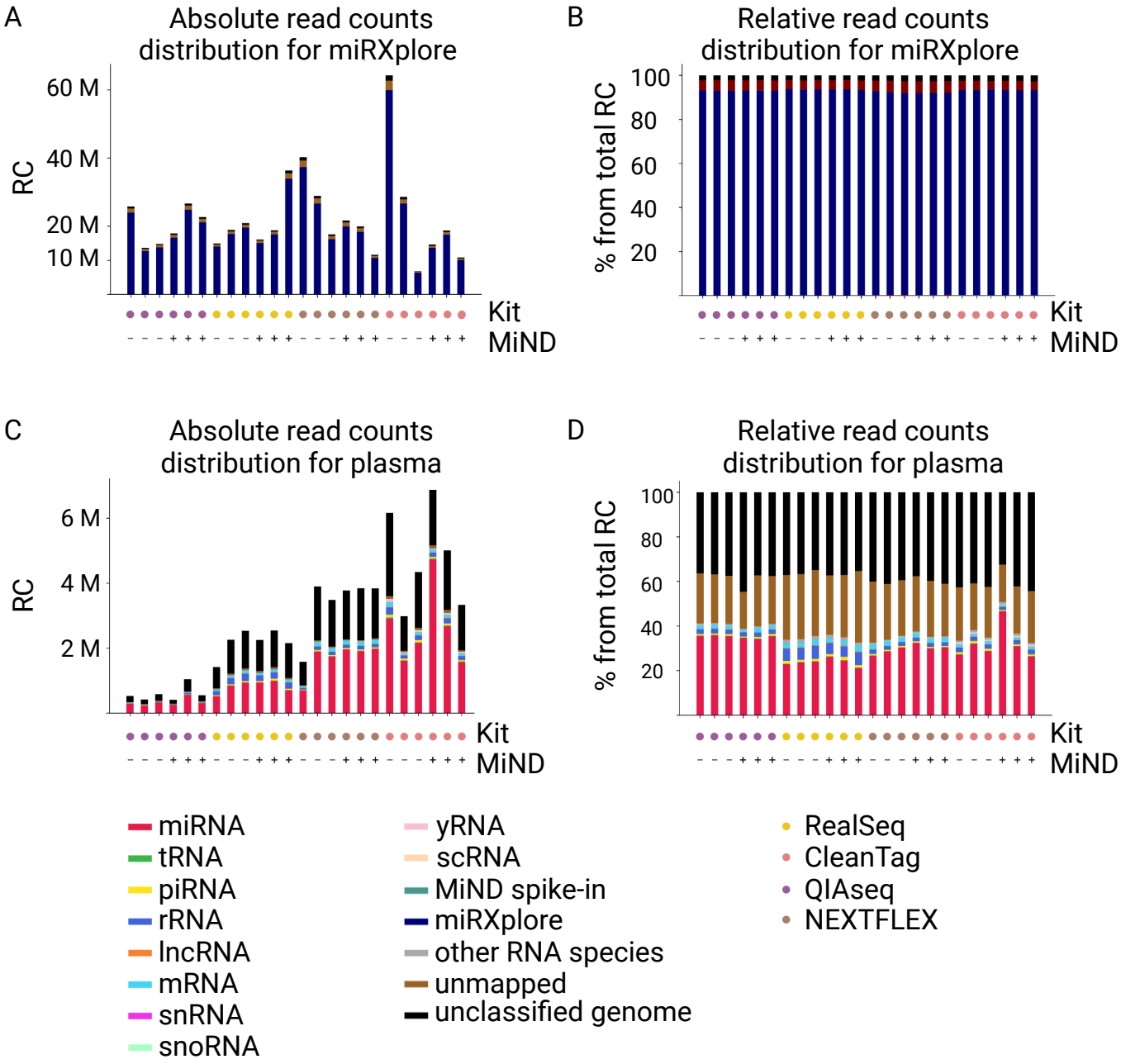


Figure S2. Number of distinct microRNAs

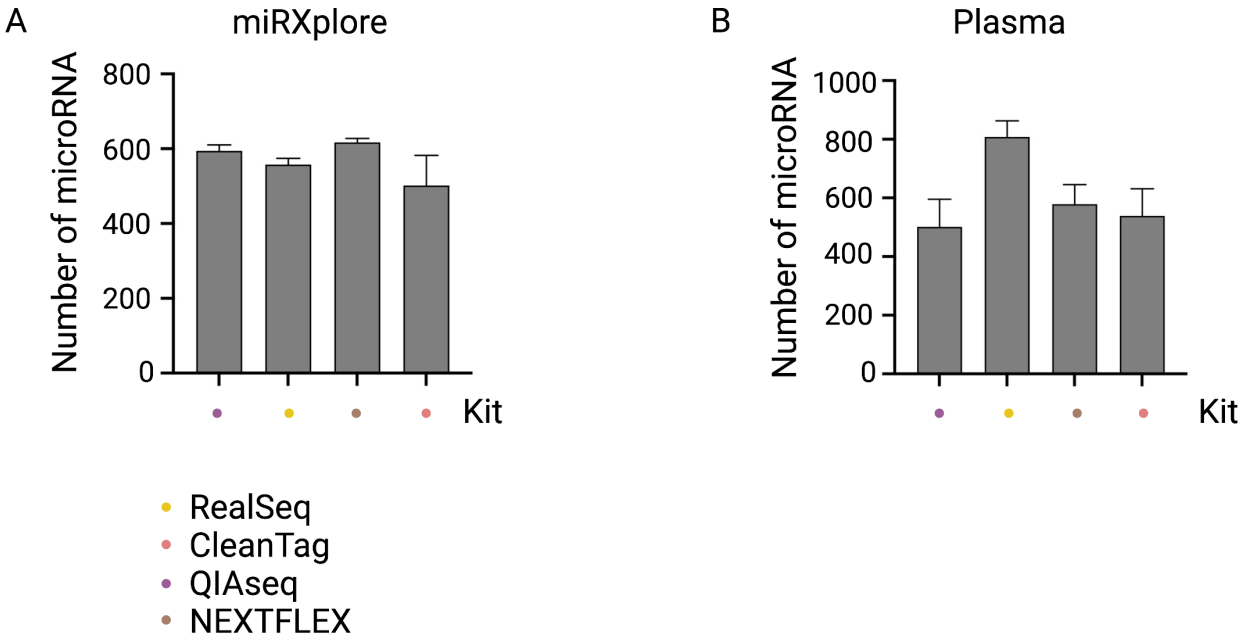


Figure S3. Performance of the selected small RNA sequencing protocols

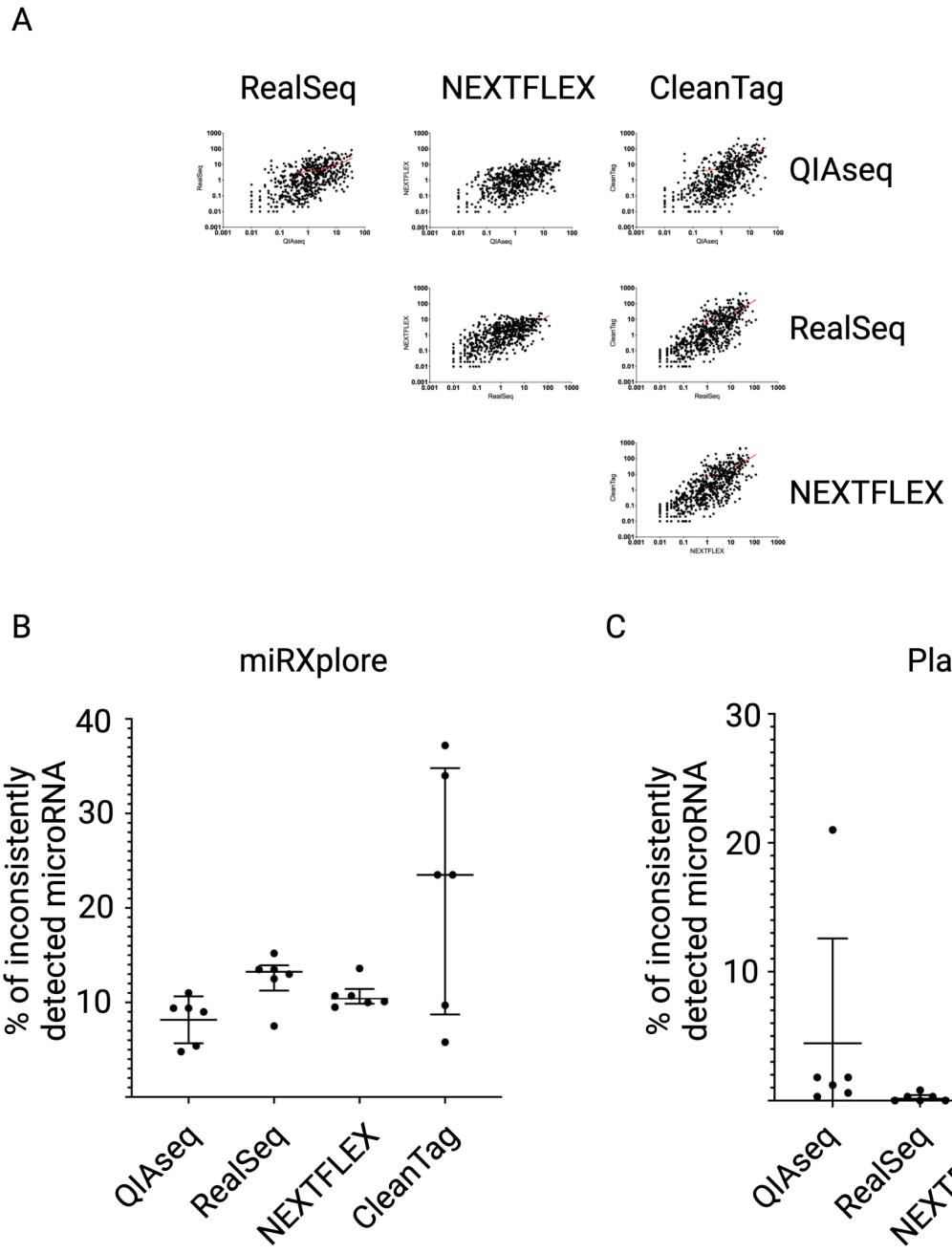
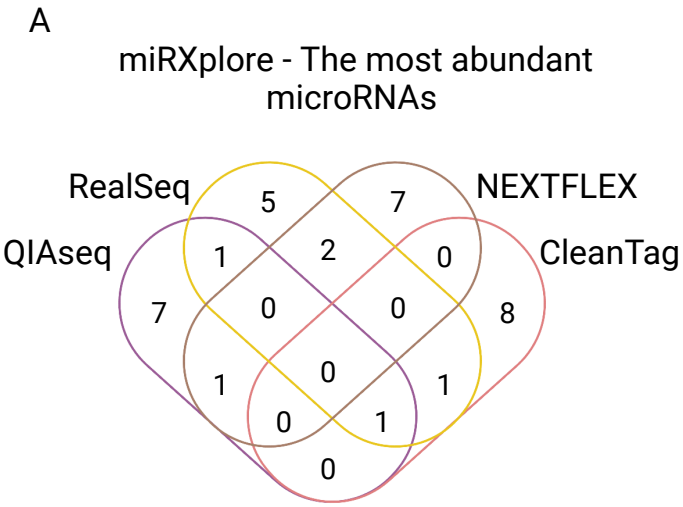
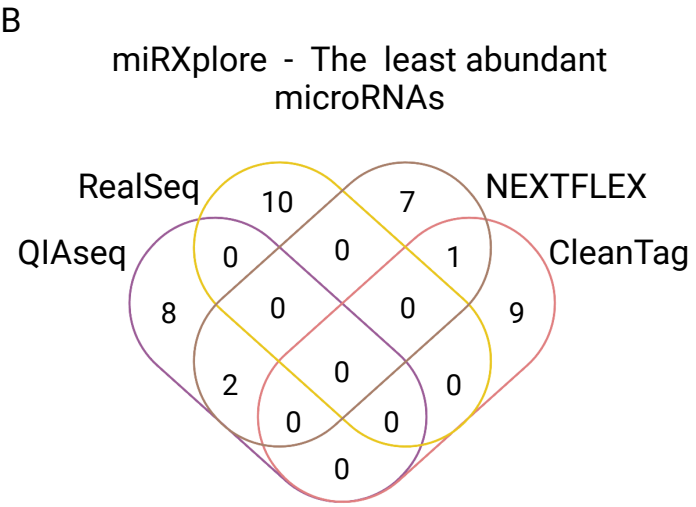


Figure S4. The 10 most and least abundant microRNAs

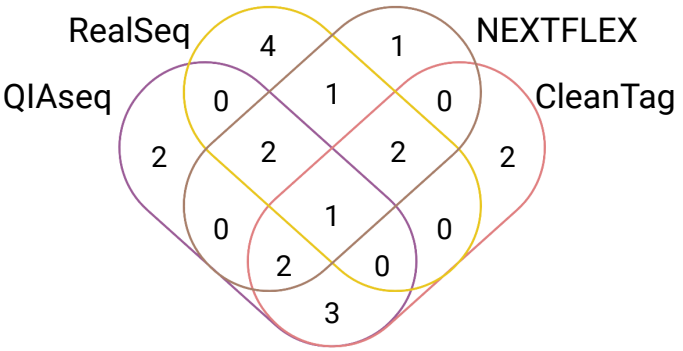
A



B



C



D

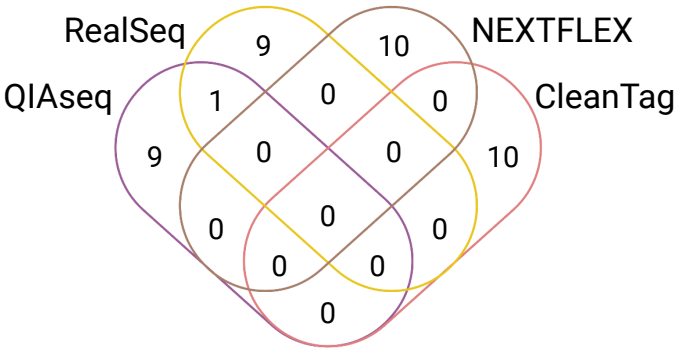


Figure S5. Scatter plots of the MiND spike-ins relative and absolute levels

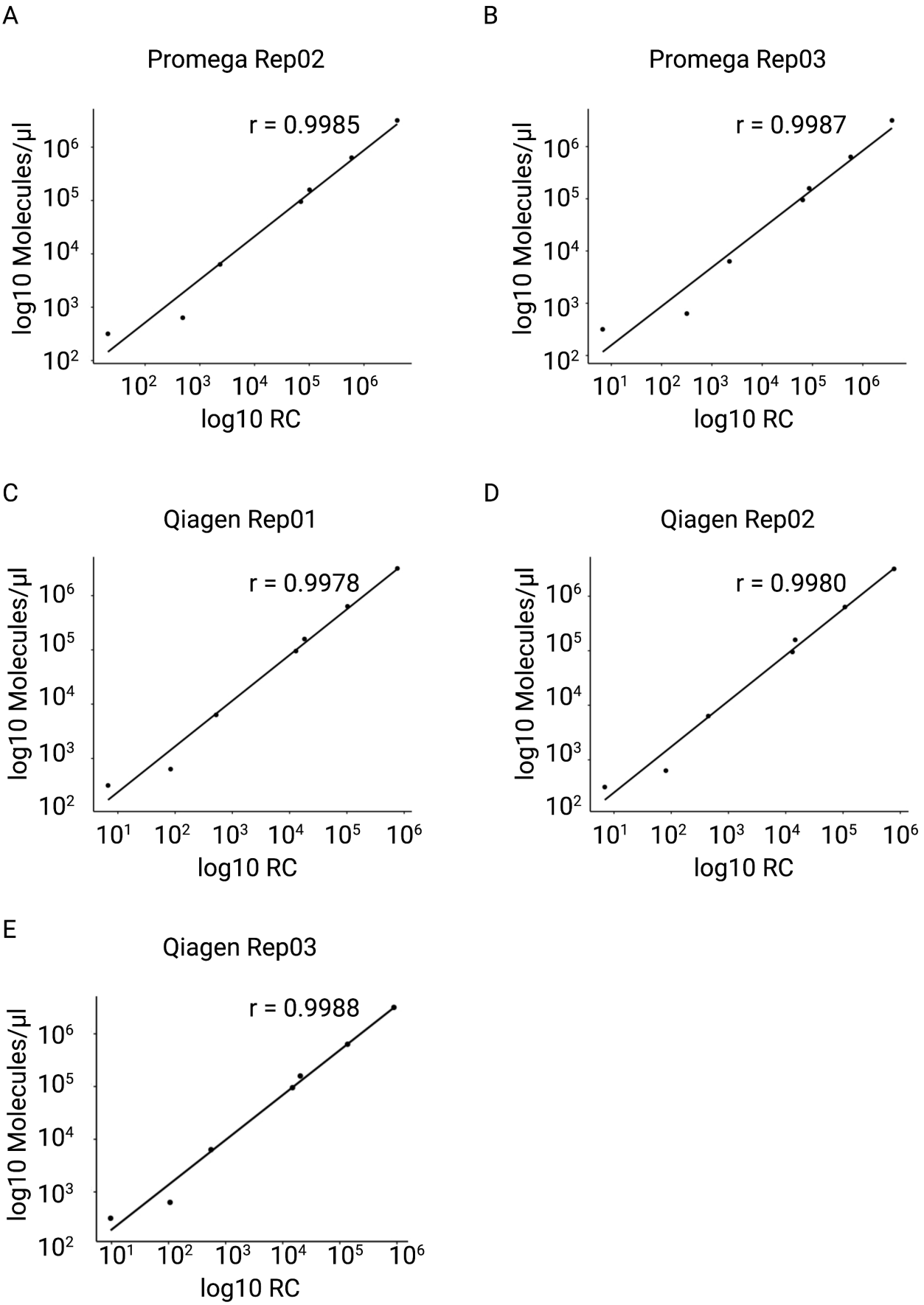


Figure S6. Optimisation of the small RNA NGS protocol

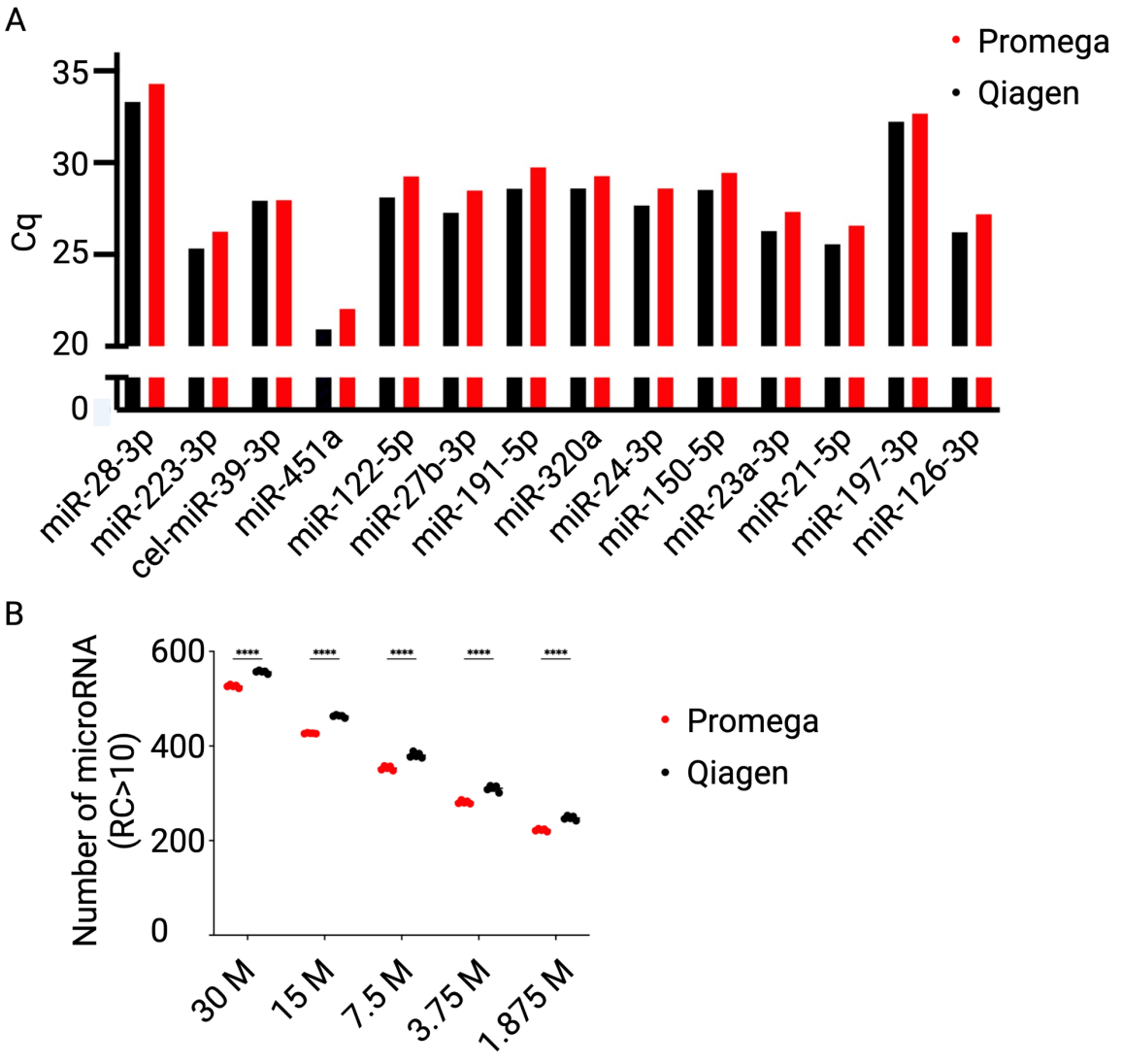


Figure S7. Impact of sequencing depth on number of detected microRNAs

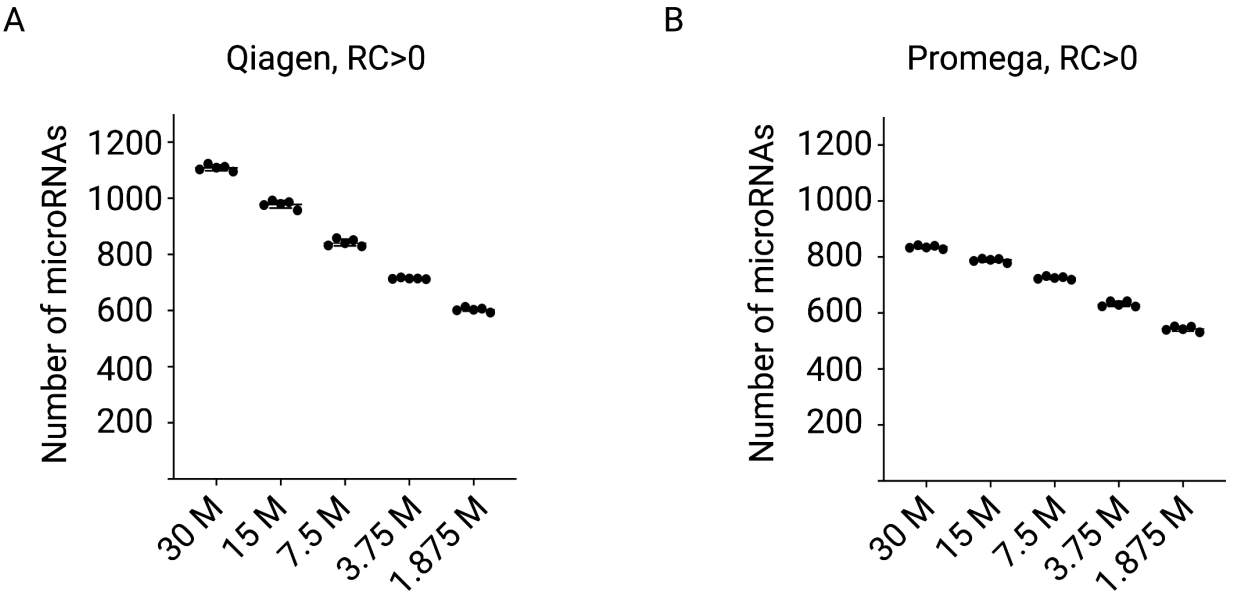


Figure S8. The MiND assay validation

