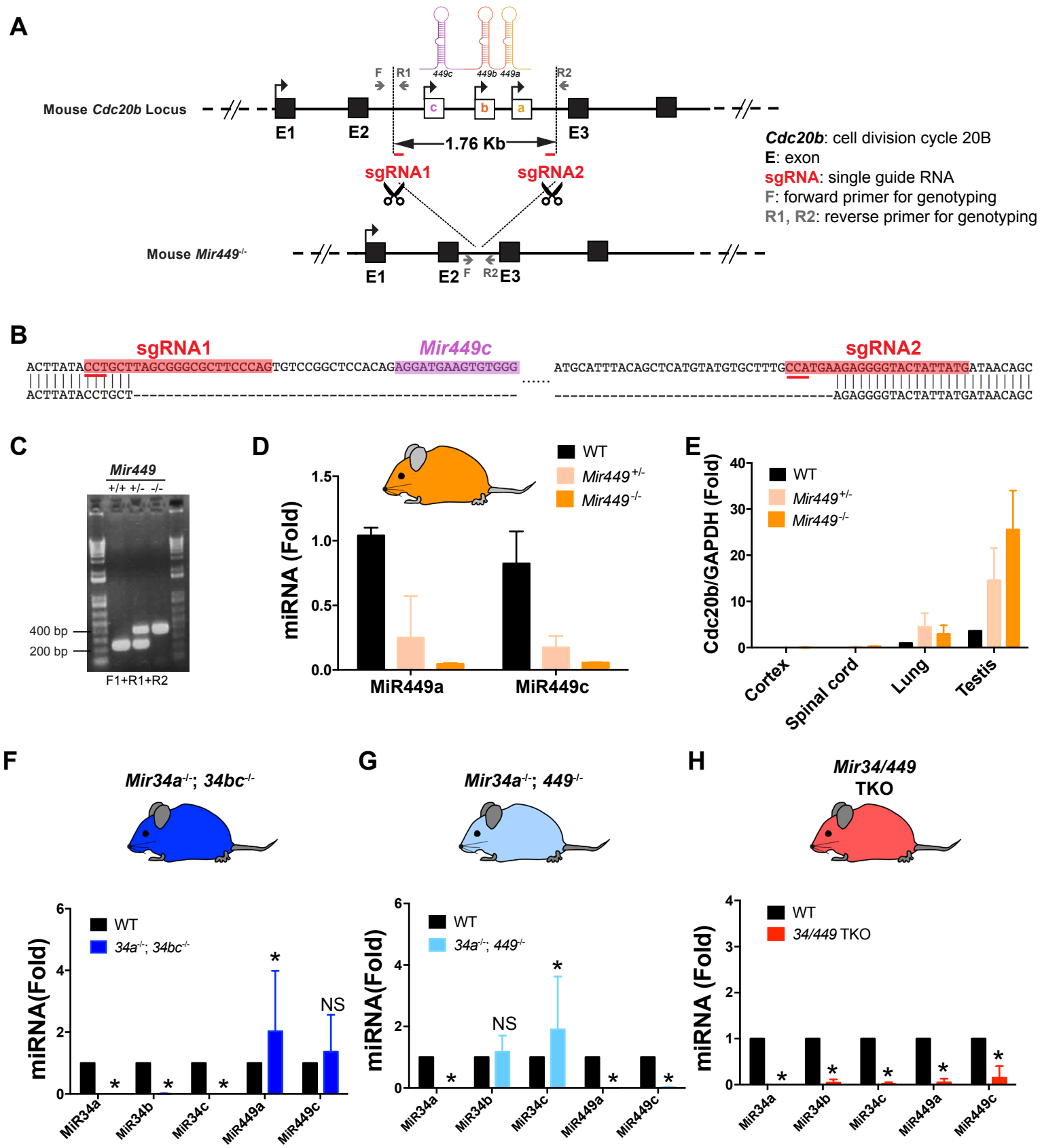Figure 1—figure supplement 1