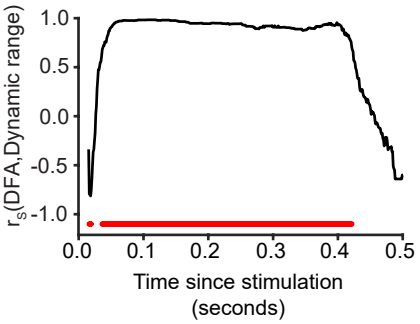Figure 1—figure supplement 2