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The influence of different Stop-signal response time estimation procedures on behavior-behavior and brain-behavior correlations

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Abstract

The fundamental cognitive-control function of inhibitory control over motor behavior has been extensively investigated using the Stop-signal task. The critical behavioral parameter describing stopping efficacy is the Stop-signal response time (SSRT), and correlations with estimates of this parameter are commonly used to establish that other variables (e.g., other behavioral measures or brain activity measures) are closely related to inhibitory motor control. Recently, however, it has been argued that SSRT estimates can be strongly distorted if participants strategically slow down their responses over the course of the experiment, resulting in the SSRT no longer reliably representing response-inhibition efficacy. Here, we performed new analyses on behavioral and functional data from an fMRI version of the Stop-signal task to gauge the consequences of using different SSRT estimation approaches that are differentially prone to the influence of strategic response slowing. The results indicate that the SSRT estimation approach can dramatically change behavior-behavior correlations. Specifically, a correlation between the SSRT and Go-trial accuracy that was highly significant with one estimation approach, virtually disappeared for the other. Additional analyses indeed supported that this effect was related to strategic response slowing. Concerning brain-behavior correlations, only the left anterior insula was found to be significantly correlated with the SSRT within the set of areas tested here. Interestingly, this brain-behavior correlation differed little for the different SSRT-estimation procedures. In sum, the current results highlight that different SSRT-estimation procedures can strongly influence the distribution of SSRT values across subjects, which in turn can ramify into correlational analyses with other parameters.

Keywords

inhibitory motor control; Stop-signal task; Stop-signal response time; brain-behavior correlations; human

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1. Introduction

The ability to suppress unwanted behavioral responses is critical for successful goal-directed behavior, as it allows individuals to quickly adjust to a changing environment by overcoming pre-potent responses when they are inadequate or inappropriate (see e.g., [1] for a review). Interest in this topic has dramatically increased over the past several years in accord with the fundamental role of inhibitory functions in normal human behavior and development, as well as in a variety of neurological and psychiatric conditions, including attention-deficit hyperactivity disorder (ADHD) and substance abuse [2–6].

Typically, these inhibitory functions are assessed by means of versions of the Go-NoGo or Stop-signal tasks. The Stop-signal task in particular has received a lot of attention recently, partly because it is thought to be relatively robust against strategic adjustments by the participants. In this task a choice-reaction Go-stimulus is, on a minority of trials, rapidly followed by a Stop-stimulus that requires the participant to cancel the response to the Go-stimulus [7, 8]. A wide range of neuroscientific approaches have yielded converging evidence that a mostly right-hemisphere network of brain areas is critical for response inhibition (but see [9]). This network includes the inferior frontal gyrus (IFG; particularly the frontal operculum extending into the insula) and the pre-supplementary motor area (pre-SMA) which are thought to interact with the basal ganglia, and the thalamus to inhibit a motor response (for reviews see [4, 5, 10, 11]).

One of the key advantages of the Stop-signal task is its strong foundation in formal computational modeling [5, 7, 12, 13]. Such explicit modeling permits the estimation of parameters that cannot be observed directly, such as the Stop-signal response time (SSRT), which estimates how quickly participants can cancel an already-initiated response (usually around 200 ms [7, 14]). Because this parameter has to be estimated from the distributions of Go-trial response times and the Go-Stimulus--Stop-Stimulus stimulus onset asynchronies (Go-Stop SOAs) on Stop-trials, a variety of estimation procedures have been developed (for reviews see [15, 16]). In one commonly used approach, a tracking procedure is used during the experiment to dynamically adjust the Go-Stop SOA to yield 50% successful Stop-trials. Under such conditions the ‘mean approach’ can be used, which derives the SSRT by subtracting the mean Go-Stop SOA from the mean Go-trial RT (henceforth termed $SSRT_m$ here). An alternative approach is the so-called ‘integration approach’. In contrast to the mean approach, the integration approach can be used, not only in the presence of a successful tracking procedure, but also if the stopping-success rate is not 50%. To achieve this flexibility, the Go-trial RTs are rank-ordered, and the RT value at the percentile that corresponds to the percentage of failed inhibitions is determined on a per-subject basis (e.g., the RT at the 61st percentile of the Go-RT distribution for a participant with 61% unsuccessful Stop-trials). The SSRT is then estimated by subtracting the average Go-Stop SOA from this RT value (henceforth termed $SSRT_i$). Alternatively, the corresponding RT percentile can be determined for all (or for the central) Go-Stop SOAs and the respective results are averaged. Nonetheless, as the real SSRT is unknown, and both of the above approaches (and others) are only estimating procedures, it remains an open question as to which approach should be used.¹

Notably, it has recently been argued that strategic/motivational aspects of performing the Stop-signal task can systematically distort the estimate of the SSRT [17], in particular if they result in more than 50% successful Stop-trials. Such a deviation from an even ratio can result from strategic response slowing throughout the experiment (i.e., when participants try to maximize their number of successful Stop-trials by ignoring the instruction to always respond quickly to the Go-stimuli), even in the presence of an adaptive procedure that

adjusts the Go-Stop SOA based on the participants' performance. Considering that such slowing can easily occur in these sorts of studies, this potential SSRT-estimation inaccuracy is important to take into account. From a theoretical perspective, the SSRT_i estimate should be more robust against such influences than simple versions of the SSRT_m because it takes into account variations in the ratio of successful Stop-trials. Here we perform new analyses on data from a recent study in which we used an adapted version of the Stop-signal task that, as a sensory control condition, included task-blocks where the Stop-stimuli were irrelevant to the task (Fig. 1; [18]). In our earlier report, we used the mean approach to estimate the SSRT (SSRT_m) and found a strong negative correlation with general task accuracy (most importantly with Go-trial accuracy), indicating that participants with high Go-trial accuracy were also better inhibitors (i.e., had shorter SSRTs). Interestingly, SSRT_m and task accuracy both correlated with activity estimates in the left anterior insula.

Based on the concern that strategic slowing might have affected our results, we wanted to directly assess the influence of SSRT-estimation procedures on across-subject correlational analyses between this parameter and others, including behavioral parameters that estimate the tendency of participants to strategically slow their Go-trial responses in a Stop-signal task. Beyond general differences in behavior-behavior and brain-behavior correlations between the SSRT_m and the SSRT_i, we were particularly interested in whether the aforementioned correlation between SSRT_m and task accuracy in this dataset [18] might have been related to the influence of strategic response slowing on the SSRT_m estimate. Specifically, if different degrees of response slowing biased our SSRT_m estimates, it would be conceivable that this same response slowing could have also ramified into higher Go-trial accuracy. Such an influence on both variables might thus explain the strong correlation between the two, in that they would both be attributable to strategic slowing. If that were the case, the additional question would arise whether the brain-behavior correlation between insula activity and SSRT_m that we also reported in this earlier publication would be affected in a similar fashion.

2. Experimental procedure

2.1. Participants

Eighteen participants took part in this study, of whom two had to be excluded due to technical problems, and another one due to poor behavioral performance, thereby resulting in fifteen participants that were included in the analyses (9 female; mean age 22.9 years). All participants had normal or corrected-to-normal visual acuity, and did not report a history of psychiatric or neurological disorders. Written informed consent was obtained before the experiment in accordance with the Duke Institutional Review Board and participants were compensated for their participation at the rate of \$20 per hour.

2.2. Task

The present experiment entailed two variants of the typical Stop-signal task [14]. Stop-relevant blocks represented the typical Stop-signal task in that participants were instructed to try to withhold their response to the Go-stimulus when a Stop-stimulus occurred, whereas in

¹In order to investigate this heterogeneity, we have looked at the sample of Stop-signal tasks in a recent meta analysis that reviewed functional-imaging studies on that topic [11]. The meta analysis contained 21 studies that employed the Stop-signal task. Of those, three did not report an SSRT estimate. Further idiosyncrasies aside, eleven of the remaining studies used the average of the Go-Stop SOA values for the calculation, of which four used the mean approach, four the integration approach (note that three of those studies did not use an adaptive staircase procedure), and three more the median approach (the same as the mean approach but using the median of the Go-RT distribution instead). The remaining seven studies used variants of the mean or median approach but did not use the full range of Go-Stop SOA values for the estimation (i.e., these studies either waited for the Go-Stop delay staircase to stabilize or tried to estimate the specific Go-Stop SOA at which the probability of a successful inhibition was exactly 50%).

Stop-irrelevant blocks the visual stimulation was identical, but participants were instructed to ignore the Stop-stimuli (i.e., to always respond; [18, 19]). Each task was performed once per experimental run (approximately 2.5 minutes each per run), separated by a 16-sec break. Odd runs began with the Stop-relevant task, even runs with the Stop-irrelevant task. Ten runs were collected for each participant, yielding a total of 943 trials per participant. The trial onset asynchrony was varied pseudo-randomly between 2 and 8 seconds (gamma distribution; average 3.2 sec) to allow for the separation of different conditions in an event-related fMRI analysis [20].

Stop-relevant blocks used a standard Stop-signal task (Fig. 1A) with a random sequence of Go-trials (80% of trials) and Stop-trials (20% of trials). On Go-trials, the orientation of a Go-stimulus had to be discriminated (oriented to the left: right index finger; oriented to the right: right middle finger). On Stop-trials, the Go-stimulus was quickly followed by a Stop-stimulus, indicating that the response to the Go-stimulus was to be canceled, yielding successful (SST) and unsuccessful Stop-trials (UST). Total stimulus duration was always 800 ms. During Stop-trials, the Go-Stop SOA was titrated using an adaptive staircase procedure to yield similar numbers of SST and UST for each participant. The staircase procedure increased the SOA by 17 ms after SST and decreased it by the same amount after UST (initial SOA: 200 ms).

Visual stimulation during Stop-irrelevant blocks was identical to that of Stop-relevant ones (Fig. 1B), but participants were instructed to respond to all Go-stimuli irrespective of the occurrence of Stop-stimuli. To maximally equate the sensory stimulation between the two block types, we also varied the Go-Stop SOA in Stop-irrelevant blocks. Specifically, SOA values were varied in a random one-up/one-down fashion after each Stop-irrelevant Stop-trial within \pm three 17-ms steps of the initial value. All staircases used the end value of the preceding Stop-relevant block as the start value.

2.3. Behavioral analysis

The key parameter describing stopping efficacy that is typically derived in a Stop-signal task is the SSRT. It describes by how much the presentation of the Stop-stimulus needs to precede the moment when a response would be executed so that this response can still be canceled, i.e. how much time the brain needs to implement response cancellation. Here, we used two different procedures to estimate this parameter: the *mean approach* (yielding what we term the SSRT_m) derives the SSRT by subtracting the mean Go-Stop SOA from the average Go-trial response time [8]. This procedure has been argued to be the most reliable approach for estimating the SSRT, if (and only if) the ratio of successful and unsuccessful Stop-trials is 50% [16]. However, small deviations from this even ratio can occur even in the presence of a tracking procedure that adjusts the Go-Stop SOA to different Go-trial response speeds. To directly address the implication of this potential issue here, we additionally estimated the SSRT using a version of another common approach, the *integration approach* [14], yielding an estimate we label SSRT_i. Here, Go-trial RTs are rank-ordered, and the RT value at the percentile that corresponds to the percentage of unsuccessful Stop-trials is determined on a per-subject basis (e.g., 61st percentile of the Go-RT distribution for a participant with 61% unsuccessful Stop-trials). The average Go-Stop delay is then subtracted from this Go-RT value. By taking deviations from an even ratio between successful and unsuccessful Stop-trials into account, this approach is more robust against such variations. Note, however, that other methods for estimating the SSRT exist, including some approaches that might be less prone to strategic response slowing as well. Such approaches include versions of the mean approach that aim to limit the Go-Stop SOAs to a subset of values that lead to a ~50% stopping success, rather than simply averaging the full range of Go-Stop SOAs [16].

Since strategic response slowing was of prime interest in this study, we endeavored to quantitatively characterize this parameter for each participant. Similarly to the study by Leotti and Wager [17], we derived two different parameters to describe this aspect: 1) as a measure of average response slowing due to task-context, we subtracted the average Stop-irrelevant Go-trial RT from their Stop-relevant-block counterpart. This parameter is closely related to *pro-active response slowing* (e.g., [21]; a term we will use here, which has been used to describe the tendency of subjects to generally slow down their Go-trial responses in the context of Stop-trials). 2) We computed how much participants slowed down their Stop-relevant-block Go-trial responses over the course of the experiment. Such ongoing slowing is in fact necessary to achieve a high proportion of successful Stop-trials in that it is the only way to avoid the staircase algorithm “catching up” and increasing the Go-Stop SOA sufficiently to counteract slow Go-responses. This parameter was estimated by computing the difference in Stop-relevant-block Go-trial RT from the second minus the first half of the experiment (here termed *progressive response slowing*).

Across-subject, behavior-behavior correlations were calculated between the following parameters: SSRT_i, SSRT_m, proportion of successful Stop-trials, Stop-relevant-block Go-trial accuracy, pro-active response slowing, and progressive response slowing. Statistical analyses of the behavioral data were performed using Pearson’s correlation. One-tailed p-values are reported because clear predictions could be made for the direction of the key correlations (e.g., that response slowing leads to shorter SSRT_m estimates and to more accurate Go-trial responding, etc.). In order to statistically compare them, we transformed correlation coefficients into z scores and estimated the difference between coefficients in relation to the standard error of this difference.

2.4. fMRI data acquisition and analysis

MR data was acquired on a 3-Tesla GE Signa MRI scanner. Functional images were acquired with a reverse-spiral sequence (TR=2000 ms, TE=25 ms; flip angle =75°; 32 slices with 3×3×3 mm resolution; AC-PC orientation covering the brain from the top approximately down to the Pons). To reach steady-state magnetization, the first five functional images of each run were discarded. Additionally, a high-resolution structural T1 (3D Fast Spoiled Gradient Recalled (FSPGR); 1×1×1 mm resolution) was obtained. Data were analyzed using SPM5 (<http://www.fil.ion.ucl.ac.uk/spm/>) including the following steps: correction for acquisition time delay; spatial realignment; spatial normalization (applying the normalization matrix used to warp the individual anatomical images onto the SPM template); reslicing to a voxel size of 2×2×2 mm; and spatial smoothing (8-mm full-width half-maximum Gaussian kernel). First-level statistical modeling was based on canonical hemodynamic response functions (HRF) combined with time and dispersion derivatives, while additionally applying a 128-sec high-pass filter [22]. All major conditions were modeled separately and restricted on trials with correct responses (or no response for successful Stop-trials). Additional regressors modeled trials with incorrect responses, misses, the onsets of breaks, as well as the six spatial realignment parameters.

The parameter estimates resulting from each condition/contrast and participant (first-level analysis) were entered into a second-level, random-effects group analysis using one-sample t-tests. For visualization purposes, activation maps were rendered on the SPM single-subject template. Since this report is a follow-up on our earlier work and is based on portions of the same dataset [18], the only brain-activity contrast used here for correlations with behavioral parameters is the between-block conjunction that was used for brain-behavior correlations in that earlier report. Specifically, a conjunction of successful and unsuccessful Stop-trials respectively compared against Stop-irrelevant Stop-trials (i.e., a sensory control without response inhibition) was performed and thresholded at $p < 0.002$ with an extent threshold $k = 10$ contiguous voxels ($p < 0.005$ and $k = 5$ for subcortical activations; see Tab. 5 in [18]).

The resulting local maxima were used for a region-of-interest (ROI) analysis, in which Marsbar (<http://marsbar.sourceforge.net/>) was used to extract beta weights from spherical 4-mm radius ROIs centered on those local maxima (see Tab. 5 in [18] for the full list of ROIs). As in our previous report, we extracted the differences of the beta values corresponding to the contrasts “SST vs. Stop-irrelevant Stop-trials” and “UST vs. Stop-irrelevant Stop-trials”, and these two values were then averaged for each subject to capture the degree to which activity in these ROIs was more pronounced during Stop-relevant vs. Stop-irrelevant Stop-trials.

In order to probe brain-behavior relationships between activity levels in the different ROIs and the SSRT measures, we correlated the activity estimates extracted from the ROIs with the SSRT_m, Go-trial accuracy (already reported in [18]), and SSRT_i values across participants. One-tailed p-values are reported for these correlational analyses, in that we expected negative correlations with the SSRT estimates in areas related to stopping efficiency. Additional brain-behavior correlations based on the same data can be found in our earlier report [18].

3. Results

3.1. Behavioral data

Table 1 provides an overview of the response-time and accuracy parameters. During Stop-relevant Stop-trials, participants managed to withhold their behavioral response on approximately half of the trials, indicating that our staircase SOA-adjustment procedure had been relatively successful in this regard. Importantly, these values varied quite closely around 50% ranging between 46% and 59% successful Stop-trials. The SSRT estimated by the two procedures yielded very similar average values. The mean approach (SSRT_m) yielded 230 ms, whereas the integration approach (SSRT_i) resulted in 225 ms, which did not differ significantly (two-tailed paired t-test: $p > 0.3$). Nevertheless, the two estimates appeared to differ somewhat within participants, as evidenced by a correlation coefficient of only 0.67 of the two estimates across participants (Fig. 2A). While highly significant ($p = 0.003$), this correlation is somewhat modest given that both variables estimate the same parameter and result in highly similar grand-average values.

As reported in our previous article using the mean SSRT-estimation approach only [18], the SSRT_m values correlated inversely with task accuracy in Stop-relevant-block Go-trials to a very high degree ($r = -0.81$, $p < 0.001$). In contrast, the SSRT_i values calculated here produced a considerably different result (Fig. 2B), in that it did not yield a significant correlation with task accuracy ($r = -0.33$; $p > 0.1$; see Tab. 2 for an overview of behavior-behavior correlations). Importantly, the two correlation coefficients differed significantly from one another (one-tailed $p = 0.001$). One possible explanation for why the SSRT_m correlates with Go-trial accuracy (and the SSRT_i does not) could be related to the influence of strategic slowing on the SSRT_m. Leotti and Wager [17] report that SSRT estimates are shorter for subjects that slow down their Go-responses over the course of the experiment in order to achieve more than 50% successful Stop-trials. Responding more slowly, in turn, is likely to also lead to higher Go-trial accuracy. Since the SSRT_i estimate is less susceptible to such influences, the observation that it is not correlated with Go-trial accuracy is consistent with this notion.

To further investigate whether strategic slowing contributed to the correlation pattern described above, we performed additional analyses based on the two parameters we derived to describe strategic slowing (see section 2.3). On average, participants slowed down their response by 84 ms during Stop-relevant blocks as compared to Stop-irrelevant blocks (*proactive response slowing*). Additionally, we calculated whether subjects slowed down their

Go-trial responses in Stop-relevant blocks over the course of the experiment by comparing Stop-relevant-block Go-trial RTs from the second minus the first half of the experiment (*progressive response slowing*). Although this value was not significantly different from zero across subjects (two-tailed one-sample t-test: $p > 0.4$), there was a large degree of variation such that some subjects sped up while others slowed down their responses over the duration of the experiment.

Performing additional correlational analyses with these two indices of strategic slowing yielded some evidence in favor of the notion that strategic slowing (and the resulting higher proportion of successful Stop-trials) had some influence on the SSRT_m, but not on the SSRT_i. More specifically, the SSRT_m was correlated with the proportion of successful Stop-trials ($r = -0.49$; $p = 0.03$), and there was a trend for a correlation with progressive response slowing ($r = -0.38$; $p = 0.08$). Neither correlation was present for the SSRT_i (both $p > 0.25$). In particular, the correlations with the proportion of successful Stop-trials yielded significantly different correlation coefficients for SSRT_m and SSRT_i (one-tailed $p = 0.001$). Moreover, the correlation coefficients for progressive response slowing also differed significantly for the correlations with SSRT_m and SSRT_i (one-tailed $p = 0.02$). Importantly, Stop-relevant Go-trial task accuracy was correlated with progressive response slowing ($r = 0.62$; $p = 0.007$) and with the proportion of successful Stop-trials ($r = 0.64$; $p = 0.005$). Additionally, there was a trend towards a significant correlation between Stop-relevant Go-trial accuracy and pro-active response slowing ($r = 0.4$; $p = 0.07$). Finally, participants who had a higher proportion of successful Stop-trials displayed more pro-active response slowing ($r = 0.84$; $p < 0.001$) and more progressive response slowing ($r = 0.66$; $p = 0.004$), whereas none of the other correlations among this set of variables reached significance (Table 2).

In light of this overall set of correlational results with the two SSRT measures, a consistent pattern appears to emerge. Specifically, the strategic slowing on Go-trial responses is related to a higher proportion of successful Stop-trials, higher Go-trial accuracy, and a faster SSRT_m estimate. In contrast, the SSRT_i estimate was not significantly correlated with either of these parameters, presumably indicating that it is indeed more robust against subtle influences of strategic response slowing.

3.2. Brain-Behavior correlation

Figure 3 provides an overlay of the grand-average results for the conjunction contrast of successful and unsuccessful Stop-trials vs. Stop-irrelevant Stop-trials. The activated areas isolated in this contrast included, among some others, the left and right IFG protruding into the insula, the pre-SMA, the caudate nucleus, and the thalamus (see Tab. 5 in [18] for the full list of 13 local activation maxima). As detailed in our previous report on this dataset, only a local maxima in the left anterior insula correlated with the SSRT_m ($r = -0.69$; $p = 0.002$), which also correlated with Stop-relevant Go-trial accuracy ($r = 0.66$; $p = 0.004$).

The pattern of results of our present reanalysis of this brain-behavior data contrasts with the dramatic effects on behavior-behavior correlations. Specifically, the correlation between brain activity in the left anterior insula and stopping efficacy that we reported before was still significant when using the SSRT_i ($r = -0.58$; $p = 0.011$), and the correlation coefficients did not differ significantly between the SSRT_i and the SSRT_m (two-tailed $p = 0.55$). For the SSRT_i, the correlation with the local maximum in the left thalamus also reached significance, but only if not corrected for multiple comparisons ($r = -0.52$; $p = 0.047$), whereas no other areas displayed correlations with the SSRT_i (all $p > 0.05$). Thus, among the set of activated areas in the investigated contrast, the SSRT estimation procedure appeared to have little influence on correlations between brain activity and SSRT. It is important to point out, however, that the present analysis was based on a relatively specific contrast that identified

only a small set of areas, so that our conclusions concerning brain-behavior correlations with SSRT_i and SSRT_m are limited to these areas.²

4. Discussion

The present study investigated the influence of two different procedures of estimating stopping efficacy (i.e., the SSRT) in the Stop-signal task on behavior-behavior and brain-behavior correlations. We found that two widely-used estimation procedures strongly influenced behavior-behavior correlations despite the fact that their grand-average values across subjects were nearly identical. In our previous report derived from the present data set, we reported that the SSRT and task-accuracy measures were strongly inversely correlated [18]. Moreover, we had found that the SSRT and task-accuracy parameters correlated with activity in the left anterior insula. These observations appeared to suggest that the SSRT did not specifically index different degrees of inhibitory ability, as previously thought, but might rather relate to more general factors such as motivation [17, 23]. Our previous analyses, however, were based on the SSRT_m estimate, which generally requires an even ratio of successful and unsuccessful Stop-trials for each participant [15, 16], a criterion that was not precisely met for all individual subjects. As noted earlier, it has been demonstrated that deviations from this ratio, which can arise from the tendency of participants to strategically slow down responses over the course of the experiment, can substantially affect the estimate of the SSRT [17].

Although in the current study the ratio between successful and unsuccessful Stop-trials was very close to 50/50 averaged across subjects, it ranged between 46% and 59% for the successful Stop-trials, varying somewhat across participants. It is therefore conceivable that the use of the SSRT_m estimate inflated the correlation with Go-trial accuracy that we have recently reported [18], in that response-slowing could have simultaneously led to shorter SSRT_m estimates and to higher Go-trial accuracy. To probe this possibility here, we recomputed the SSRT using the integration approach (SSRT_i), which is less susceptible to deviations from exactly even ratios of successful and unsuccessful Stop-trials. Although the *average* SSRT estimates across subjects did not differ significantly between the two procedures (SSRT_m: 230 ms; SSRT_i: 225 ms), the correlation pattern with other parameters differed quite substantially. In particular, the correlation with Go-trial task accuracy was indeed substantially attenuated when using the SSRT_i and was not significant, in contrast to this correlation being highly significant when using the SSRT_m. Moreover, the calculated coefficients for the correlations of accuracy and SSRT when using the SSRT_i versus the SSRT_m were significantly different from each other. Thus, it seems likely that the correlation between SSRT_m and task accuracy was indeed inflated by different degrees of strategic response slowing across participants.

Further corroboration for this notion came from additional behavior-behavior correlations. Specifically, the correlations including the ratio of successful Stop-trials, Go-trial accuracy, and the two parameters describing different aspects of strategic response slowing (pro-active and progressive) indicated that these parameters are highly correlated with one another and with the SSRT_m but not with the SSRT_i. Accordingly, it appears quite likely that progressive response slowing in particular (i.e., slowing over the course of the experiment) leads to higher rates of successful Stop-trials, higher Go-trial accuracy, and shorter SSRT_m

²Note also that the activity estimates used here are based on the average of successful and unsuccessful Stop-trials. Moreover, activity estimates for task-irrelevant Stop-trials were subtracted from both estimates. While the latter was done to enhance the specificity of the resulting values (i.e., in order to exclude activity unrelated to response inhibition), the use of both SST and UST might seem more unconventional. However, due to our use of a very small stepsize in the staircase titration procedure, it is likely that SST and UST trial types produced substantial and similar levels of stop-related activity (see [18] and Discussion below for further details). Moreover, when splitting up the brain-behavior correlations, we found that they were similar for the parameters based on SST and UST.

estimates. Taken together, these observations would argue against our previous suggestion that the SSRT might reflect more general processes rather than being specific for response inhibition abilities (see also [17, 24–27]). Our results rather appear to indicate that the SSRT, when calculated suboptimally, can be problematically influenced by other behavioral variables. In particular, it appears that individual-subject SSRT_m measures can be influenced by strategic response slowing by the participants even if it only occurs to a very moderate degree. Thus, this estimation approach should indeed only be used if the known prerequisite of an even ratio of successful and unsuccessful Stop-trials is quite precisely met. This is particularly relevant when the estimates are used for correlational analyses, because it would appear, at least in our data, that it is more the distributions of individual values that are affected, without significantly changing the average value. The SSRT_i, in contrast, appears to be relatively robust to the influence of strategic response slowing, at least when the latter only occurs to a moderate degree, in that it did not display a significant correlation with any of the RT-slowness-related parameters and task-accuracy analyzed here. Given that the SSRT_i can also be calculated quite easily, and that it generalizes to various versions of the Stop-signal task and a wider range of stopping-success rates (i.e., beyond 50%), the SSRT_i would seem to have considerable potential as a technique that can be applied in a great number of experimental contexts. However, it should be noted that other approaches exist, which may have some advantages in certain circumstances [16].

It is important to note that the influence of the SSRT estimation procedure on behavior-behavior correlations was stronger than one might have expected. In particular, the mean values across subjects from the two procedures were virtually identical, and the mean ratio between successful and unsuccessful Stop-trials was fairly close to 50% (52.7%), in which case the use of the mean approach would appear to have been warranted. Along similar lines, in the study by Leotti and Wager that investigated the influence of response slowing on SSRT estimates, the group that was defined as “non-compliant” (i.e., displaying too much response slowing) had a ratio of successful Stop-trials of 69% in their first experiment, and also the amount of RT slowing was substantial in this experiment (more than 100 ms over the course of the experiment and more than 200 ms compared to a block with speeded Go-trials only) [17]. The present set of participants was far more homogenous and displayed much less evidence of strategic response slowing, so that only subtle influences on the SSRT_m estimate would have been expected. Thus, the strong influence of the estimation approach on some of the behavior-behavior correlations was somewhat surprising.

The only brain-behavior correlation with SSRT_m we had found in our earlier report was with the left anterior insula, activity in which also correlated with general task accuracy [18]. This correlation was replicated in the present analysis for the SSRT_i, whereas no strong correlations were found in other areas for either estimate. Accordingly, in contrast to the substantial influence on behavior-behavior correlations, the left anterior insula activation does indeed appear to be related to the SSRT, independent of the SSRT estimation procedure. The additional correlation of activity in this area with task accuracy that we reported previously, however, still suggests a relatively unspecific role for this area in enabling good task performance that is not specific for response inhibition (see also [11]).

Interestingly, the left anterior insula is not commonly identified in brain-behavior correlations with the SSRT. More generally, it can be noted that in the various studies that have investigated such brain-behavior correlations, the areas in which brain activity has been reported to correlate significantly with the SSRT have varied greatly, with limited overlap across the studies [28–37]. This variation might be related to power issues and the high dimensionality of the employed brain measures [38]. Additionally, other design factors or the use of different functional contrasts for quantifying activity related to response inhibition might have played a role in producing such variation. For example, the present study used

activity estimates from a conjunction of successful and unsuccessful Stop-trials (relative to the Stop-Irrelevant control condition), as the two conditions yielded very similar activation patterns. This latter finding could be related to a race model where inhibition gets triggered for successful and unsuccessful Stop-trials, so that activity related to response inhibition should be present in both conditions [18]. Additionally, however, the SSRT estimation approach employed might have played a role in the correlation variations across studies, in that we have shown in the present study that the estimation approach can strongly influence the across-subject distribution of SSRT values and thus the correlational analyses of these values.

Lastly, we note that although the reported brain-behavior correlation with activity in the left anterior insula was largely unaffected here, we only tested a small number of areas – i.e., only those that resulted from the highly specific contrast that we employed. Thus, we caution that our present findings indicating an estimation-approach independence of the correlation with this particular area therefore do not necessarily generalize to a wider set of brain areas. In particular, it would be conceivable that brain areas that are involved in strategic response slowing could also correlate with the SSRT_m in a fashion that would in fact be confounded by the strategic response slowing itself, which could affect not only correlational analysis, but also between-group comparisons of the SSRT if the groups differ in this regard. In this context, it is important to note, however, that there have been some suggestions that strategic response slowing could in fact shorten the “real” SSRT [39, 40]. Although such an influence would represent a violation of the independence assumption of the horse-race model [8], such effects are fully conceivable. The full set of behavior-behavior correlations, and the fact that the SSRT_i did not correlate with either of our response-slowing parameters, however, appears to indicate that significant influences of strategic slowing on the “real” SSRT were likely not present here.

5. Conclusions

The present report investigated how two different SSRT estimation procedures, both of which are fairly widely used, can influence behavior-behavior and brain-behavior correlations. A surprisingly strong effect on behavior-behavior correlations between the SSRT_m and Go-trial accuracy appeared to be mediated by strategic response slowing by the participants, which was not seen for the SSRT_i estimate. On the level of brain-behavior correlations, only the left anterior insula correlated clearly with SSRT (and, in turn, with Go-trial accuracy), and in this case the specific estimation procedure had little influence on the correlation. Taken together, these results demonstrate that correlations between the SSRT and other parameters are susceptible to strategic response slowing if the SSRT is calculated with the oft-used mean approach. This drastic influence on correlations with other parameters stands in contrast to the fact that the overall grand-average SSRT estimates across subjects did not differ significantly between the two approaches. Accordingly, the present results suggest that the distribution of individual-subject SSRT values can have a strong effect on correlational measures that might not be expected based on the grand-average estimates. Thus, it would seem that these differences are important to take into consideration in Stop-signal tasks employing correlational analyses involving the SSRT measures. In general, the SSRT_i estimate appears to be less prone to such influences and thus would seem to be preferred over the standard SSRT_m, even when the pre-requisite of having exactly 50% stopping success for every participant is only slightly missed.

Highlights

1. Different methods of estimating stopping efficacy (SSRT) yield comparable mean values

2. Yet, different estimates led to very different correlations with Go-trial accuracy
3. Conversely, left insula activity and SSRT correlated strongly for both estimates

Non-standard Abbreviations

IFG	inferior frontal gyrus
pre-SMA	pre-supplementary motor area
SSRT	Stop-signal response time
SSRTi	SSRT estimated by the integration approach
SSRTm	SSRT estimated by the mean approach
SST	successful Stop-trial
UST	unsuccessful Stop-trial

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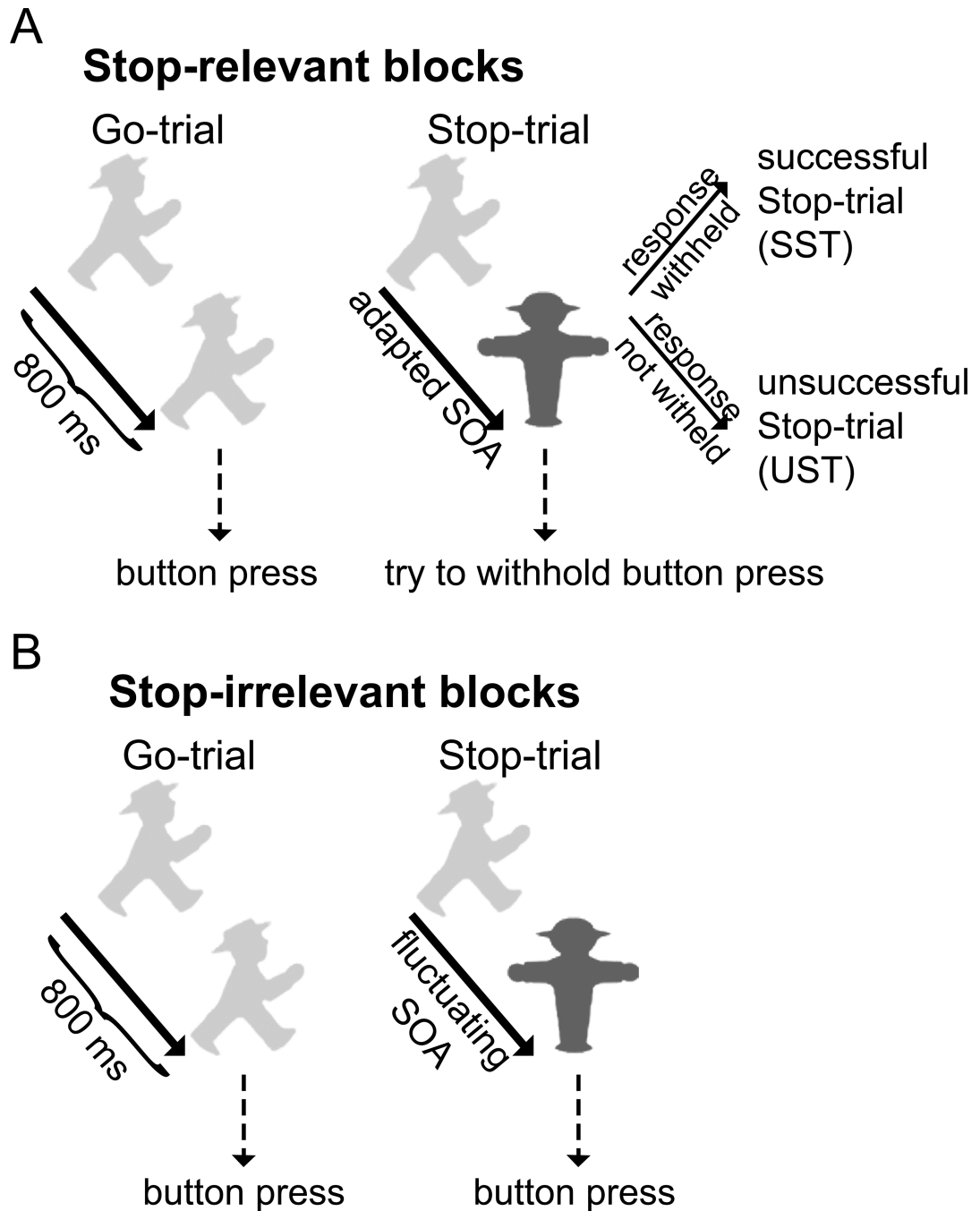


Fig. 1. Paradigm. (A) In Stop-relevant blocks, a choice-reaction stimulus (a green German traffic-light symbol oriented to the left or right – here represented in light grey – requiring an index / ring-finger response respectively) was either presented for 800 ms (Go-trial) or rapidly replaced by a red Stop-stimulus (here represented in dark grey; Stop-trial). This Stop-stimulus was presented after a variable SOA set by a tracking algorithm and indicated that the response to the Go-stimulus on that trial was to be inhibited, thereby yielding successful (SST) and unsuccessful Stop-trials (UST). (B) In Stop-irrelevant blocks the visual stimulation was identical, but the Stop-stimuli were irrelevant and behavioral responses

were required on all trials, thus providing a sensory baseline condition and a reference for response speed in the absence of task-relevant Stop-stimuli.

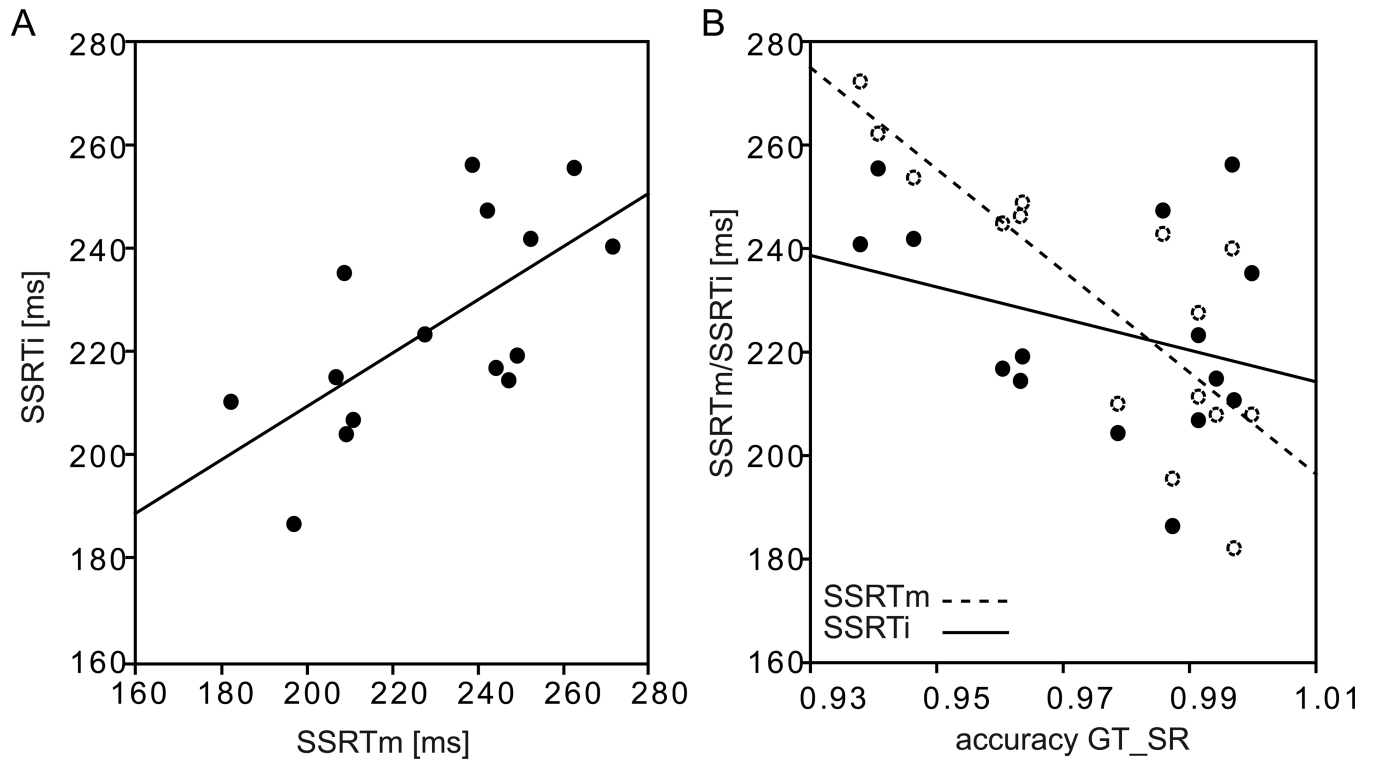


Fig. 2. Across-subject behavior-behavior correlations with two different SSRT estimates. (A) The SSRT values resulting from the two different estimation procedures were strongly correlated ($r=0.67$). (B) However, a significant correlation between SSRT and Go-trial accuracy was only found for the SSRTm estimate (SSRTm: $r=-0.81$; SSRTi: $r=-0.33$).

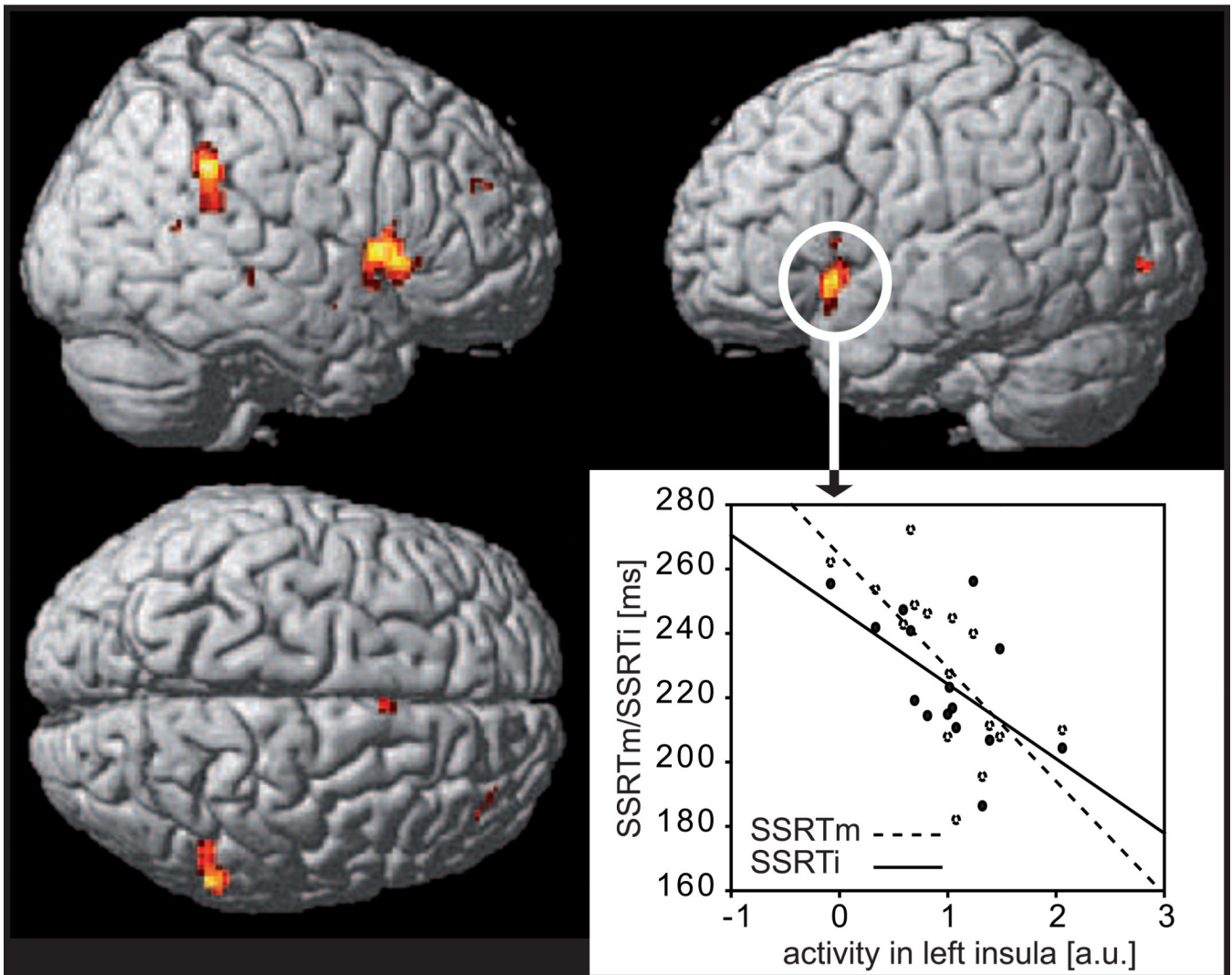


Fig. 3. Grand-average brain activity and brain-behavior correlations. The left anterior insula correlated inversely with the SSRT largely independent of the estimation method (SSRTm: $r=-0.69$; SSRTi: $r=-0.58$). This region was the only one that showed a strong relationship between brain activity and either of the SSRT measures in the present study (a.u. = arbitrary units of parameter estimates).

Table 1Behavioral data (average \pm SD)

Stop-relevant blocks	
Correct Go-trials	97.6% (\pm 2.1)
Proportion successful Stop-trials	52.7% (\pm 3.8)
RT Go-trials	520 ms (\pm 84)
RT unsuccessful Stop-trials	446 ms (\pm 64)
SSRTm	230 ms (\pm 26)
SSRTi	225 ms (\pm 20)
Progressive response slowing	4 ms (\pm 37)
Stop-irrelevant blocks	
Correct Go-trials	96.6% (\pm 2.5)
Correct Stop-trials	97.1% (\pm 3)
RT Go-trials	436 ms (\pm 48)
RT Stop-trials	439 ms (\pm 44)

SSRTm = SSRT estimate using the mean approach; SSRTi = SSRT estimate using the integration method; Progressive response slowing = Stop-relevant Go-trial RT from the 2nd half of the experiment minus that of the first.

Table 2

Behavior-behavior correlation coefficients

	SSRTm	SSRTi	Correct Stop- Relevant Go -trials	Proportion successful Stops	Progressive response slowing	Pro-active response slowing
SSRTm	-					
SSRTi	0.67 ^{***}	-				
Correct Stop-Relevant Go-trials	-0.81 ^{***}	-0.33	-			
Proportion successful Stops	-0.49 [*]	0.19	0.64 ^{***}	-		
Progressive response slowing	-0.38 ^(#)	0.09	0.62 ^{***}	0.66 ^{***}	-	
Pro-active response slowing	-0.25	0.34	0.4 ^(*)	0.84 ^{***}	0.39 ^(*)	-

SSRTm = SSRT estimate using the mean approach; SSRTi = SSRT estimate using the integration method; Progressive response slowing = Stop-relevant Go-trial RT from the 2nd half of the experiment minus that of the first. Pro-active response slowing = Stop-relevant Go-trial RT minus Stop-irrelevant Go-trial RT.

(^{*}) p<0.1;

* p<0.05;

** p<0.01;

*** p<0.001.